



**HAL**  
open science

# Multiple functionalization of C60 hexa-adducts by “click” chemistry

Franck Schillinger

► **To cite this version:**

Franck Schillinger. Multiple functionalization of C60 hexa-adducts by “click” chemistry. Other. Université de Strasbourg, 2016. English. NNT : 2016STRAF030 . tel-02003577

**HAL Id: tel-02003577**

**<https://theses.hal.science/tel-02003577>**

Submitted on 1 Feb 2019

**HAL** is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.

**ÉCOLE DOCTORALE DES SCIENCES CHIMIQUES**

**Laboratoire de Chimie des Matériaux Moléculaires – UMR 7509**

**THÈSE** présentée par :

**Franck SCHILLINGER**

soutenue le : 29 septembre 2016

pour obtenir le grade de : **Docteur de l'université de Strasbourg**

Discipline/ Spécialité : Chimie

**Fonctionnalisation Multiple d'Hexa-adduits du  
C<sub>60</sub> par Chimie « Click »**

**THÈSE dirigée par :**

**M. Jean-François NIERENGARTEN** Dr., Université de Strasbourg

**RAPPORTEURS :**

**M. Dominique LUNEAU** Pr., Université Claude Bernard de Lyon 1

**M. Fernando LANGA** Pr., Universidad de Castilla-La Mancha (Toledo)

---

**AUTRES MEMBRES DU JURY :**

**M. Michel MIESCH** Dr., Université de Strasbourg



## Remerciements.

Ce travail a été effectué au sein du Laboratoire de Chimie des Matériaux Moléculaires (Unistra/CNRS, UMR 7509) sous la direction du Dr. Jean-François Nierengarten.

J'aimerais tout d'abord remercier mon directeur de thèse, le Dr. Jean-François Nierengarten, pour m'avoir accueilli dans son groupe de recherche et pour m'avoir donné l'opportunité de réaliser ma thèse sur un sujet passionnant. Je le remercie également pour tous les conseils qu'il m'a prodigué au cours de cette thèse. Ces conseils m'ont beaucoup fait progresser sur le plan scientifique.

Je remercie chaleureusement le Dr. Michel Holler pour ses conseils et le soutien qu'il m'a apporté au cours de cette thèse. Et merci aussi pour avoir été le premier lecteur-correcteur de ce manuscrit de thèse.

Je remercie sincèrement les professeurs Dominique Luneau, Fernando Langa et Michel Miesch d'avoir accepté et pris le temps de juger ce travail de thèse.

Je tiens à remercier le Ministère de l'Enseignement Supérieur et de la Recherche pour m'avoir attribué un contrat d'allocataire de recherche pour mes trois années de thèse.

Je tiens également à remercier Dr. Thi Minh Nguyet Trinh, Dr. Sebastiano Guerra et Eric Meischner pour m'avoir aidé à réaliser une partie de ce travail de thèse, leur sympathie et leur amitié.

Je remercie Dr. Michel Schimtt pour les mesures RMN, Dr. Jean-Marc Strub pour les mesures de spectrométrie de masse et Mathieu Chessé pour les mesures analytiques diverses.

Je remercie tous les membres du laboratoire présents et passé : Uwe, Iwona, David, Bilou, Lucie, Meera, Mathilde, Haïfa, Pauline, Radian, Thibault, Xavier, Marco, Yacine, Elodie, Morgane, Eva, Thomas, Vladimir, Huixian.... pour leur bonne humeur et leur sympathie.

Un très grand MERCI à mes parents qui m'ont toujours soutenu, guidé et réconforté dans mes choix.

Enfin, je tiens à remercier toutes les personnes qui ont contribué de près ou de loin à la réalisation de mes travaux de thèse et qui n'ont pas été citées nommément



# Table of contents.

<b>Remerciements.....</b>	<b>3</b>
<b>Abbreviation.....</b>	<b>9</b>
<b>Résumé.....</b>	<b>11</b>
<b>Chapter I : General introduction. ....</b>	<b>25</b>
<b>1. Generalities.....</b>	<b>26</b>
1.1 Discovery of a new allotropic form of carbon. ....	26
1.2 Production and Purification. ....	27
1.3 Fullerene C <sub>60</sub> : Structure and Properties.....	29
1.3.1 Structure .....	29
1.3.2 Solubility .....	29
1.3.3 Photophysical properties.....	30
1.3.4 Electrochemical properties.....	31
<b>2. Reactivity of C<sub>60</sub> and its derivatives.....</b>	<b>32</b>
2.1 Introduction. ....	32
2.2 Addition and cycloaddition reactions. ....	32
2.3 Multi-functionalization of C <sub>60</sub> . ....	34
2.3.1 Nomenclature.....	34
2.3.2 Symmetry of simple multi-adducts of C <sub>60</sub> . ....	36
2.3.3 Regiochemistry of twofold additions. ....	38
2.3.4 Multiple additions on C <sub>60</sub> .....	38
<b>3. The “Click chemistry” in the fullerene chemistry. ....</b>	<b>41</b>
3.1 The “click” concept. ....	41
3.2 The <i>Huisgen type</i> 1,3-dipolar cycloaddition. ....	42

3.3 The Copper-catalyzed alkyne-azide 1,3-dipolar cycloaddition in Fullerene chemistry.....	43
3.3.1 Introduction.....	43
3.3.2 CuAAC reactions with Mono-, Bis- and Tris-adducts of C <sub>60</sub> .....	44
3.3.3 CuAAC reactions with <i>T<sub>h</sub></i> -addition pattern hexa-adducts. ....	46
<b>4. Objectives. ....</b>	<b>52</b>
<b>5. Bibliography. ....</b>	<b>53</b>
<b>Chapter II: Regioselective syntheses of C<sub>60</sub> derivatives by a macrocyclic approach. ....</b>	<b>59</b>
<b>1. Introduction. ....</b>	<b>60</b>
<b>2. Regioselective syntheses of Bis-adducts of C<sub>60</sub> by macrocyclic approach. ....</b>	<b>64</b>
2.1 Preparation of macrocyclic bis-malonates. ....	64
2.1.1 Synthesis of macrocyclic bis-malonates: the direct approach. ....	64
2.1.2 Synthesis of macrocyclic bis-malonates: the stepwise approach. ....	68
2.2 Regioselective functionalization of C <sub>60</sub> with macrocyclic bis-malonates. ....	69
2.3 Deprotection. ....	73
<b>3. Regioselective syntheses of Tris-adducts of C<sub>60</sub> by macrocyclic approach. ....</b>	<b>75</b>
3.1 Stepwise preparation of cyclo-tris-malonates.....	75
3.2 Regioselective tris-functionalization of C <sub>60</sub> .....	81
<b>4. Conclusion.....</b>	<b>83</b>
<b>5. Bibliography.....</b>	<b>83</b>
<b>6. Experimental part.....</b>	<b>84</b>

## **Chapter III: Regioselective syntheses of C<sub>60</sub> Tris-adducts by a tethered-directed approach..... 117**

<b>1. Introduction. ....</b>	<b>118</b>
<b>2. Regioselective syntheses of C<sub>60</sub> Tris-adducts.....</b>	<b>123</b>
2.1 Syntheses. ....	123
<b>3. Synthesis of optically pure [60]fullerene <i>e,e,e</i>-tris-adducts. ....</b>	<b>130</b>
3.1 Introduction. ....	130
3.2 Syntheses. ....	132
<b>4. Conclusion.....</b>	<b>135</b>
<b>5. Bibliography. ....</b>	<b>136</b>
<b>6. Experimental part.....</b>	<b>137</b>

## **Chapter IV : Syntheses of multi-functionalized hexa-adducts by sequential “click” reactions..... 153**

<b>1. Introduction. ....</b>	<b>154</b>
<b>2. Synthesis of mixed C<sub>60</sub> hexa-adducts by “click” reactions.....</b>	<b>159</b>
2.1 Preparation of mixed C <sub>60</sub> hexa-adducts building blocks.....	159
2.2 Post-functionalization of the mixed C <sub>60</sub> hexa-adducts by CuAAC reactions. ....	166
<b>3. Synthesis of mixed C<sub>60</sub> hexa-adducts by esterification and “click” reactions.....</b>	<b>172</b>
<b>4. Conclusion.....</b>	<b>176</b>
<b>5. Bibliography. ....</b>	<b>177</b>
<b>6. Experimental part.....</b>	<b>178</b>



<b>Chapter V: Elaboration of dendrimers based on C<sub>60</sub> hexa-adducts.....</b>	<b>201</b>
<b>1. Introduction. ....</b>	<b>202</b>
<b>2. Preparation of tri-, tetra- and pentafullerene derivatives. ....</b>	<b>206</b>
2.1 Preparation of the building blocks.....	206
2.2 Synthesis of tris-, tetra- and pentafullerene derivatives. ....	212
<b>3. Giant mixed C<sub>60</sub> hexa-adducts dendrimer. ....</b>	<b>217</b>
<b>4. Conclusion.....</b>	<b>221</b>
<b>5. Bibliography.....</b>	<b>221</b>
<b>6. Experimental part.....</b>	<b>222</b>
<b>Chapter VI : General conclusions. ....</b>	<b>237</b>

## Abbreviation.

br	broad
calcd.	calculated
CuAAC	Copper Catalyzed Azide-Alkyne Cycloadditions
DBU	1,8-Diazobicyclo[5,4,0]undec-7ene
DCC	N,N'-dicyclohexylcarbodiimide
DIBAL-H	Diisobutylaluminium hydride
DMAP	4-Dimethylaminopyridine
DMF	N,N'-dimethylformamide
GPC	Gel permeation chromatography
HOMO	Highest occupied molecular orbital
HPLC	High performance liquid chromatography
IR	Infrared
IUPAC	International Union of Pure and Applied Chemistry
LUMO	Lowest unoccupied molecular orbital
MALDI-TOF	Matrix Assisted Laser Desorption/Ionisation - Time Of Flight
NMR	Nuclear magnetic resonance
PMB	Para-methoxybenzyl
redox	reduction/oxydation
sh	shoulder
TBAF	Tetrabutylammonium fluoride
TFA	Trifluoroacetic acid
THF	Tetrahydrofuran
TMS	Trimethylsilyl
UV/vis.	Ultraviolet/visible
$\epsilon$	Molar attenuation coefficient



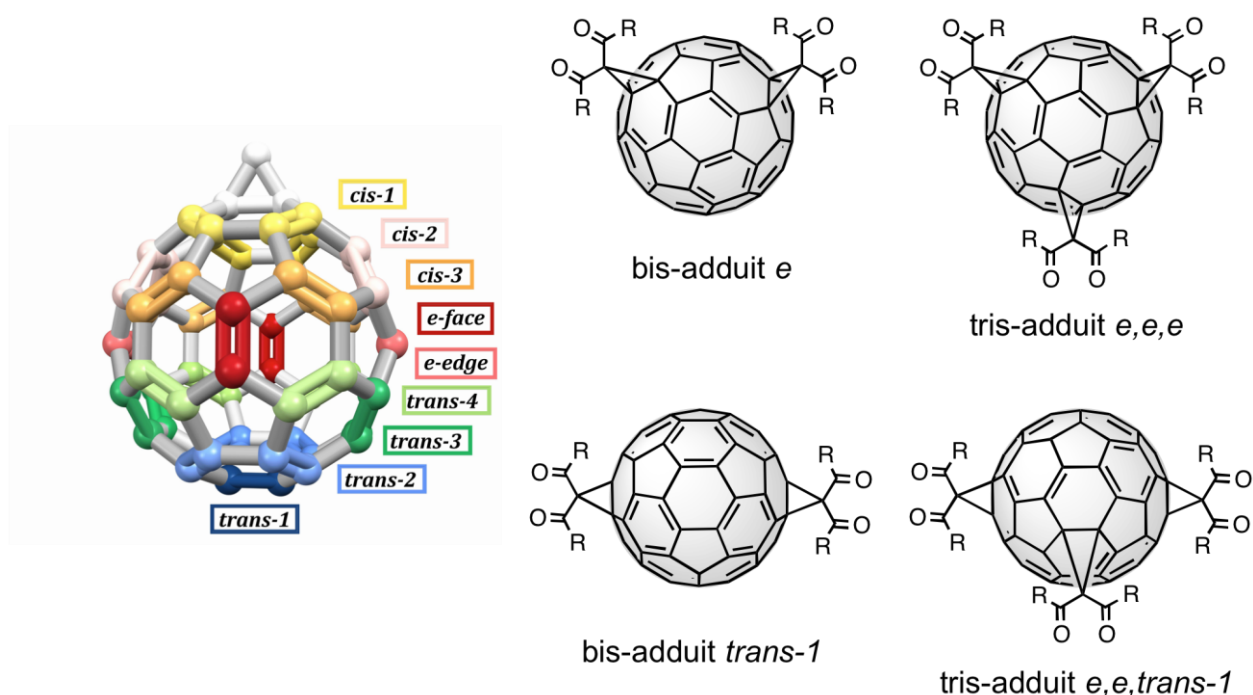
## Résumé.

Le  $C_{60}$  a été découvert en 1985 par H.W. Kroto, R.F. Curl et R.E. Smalley,<sup>[1]</sup> ce qui leur a valu le prix Nobel de chimie en 1996. Le  $C_{60}$ , aussi appelé Buckminsterfullerène, est le plus stable et le plus abondant représentant de la famille des fullerènes. Après le graphite et le diamant, les fullerènes sont la troisième forme allotropique connue du carbone. Ils correspondent à des cages creuses de carbone formés de pentagones et d'hexagones fusionnés. La chimie du  $C_{60}$  et des autres fullerènes s'est beaucoup développée dès la première production en quantité macroscopique de ces composés par W. Krätschmer et D. Huffman en 1990.<sup>[2]</sup> Les propriétés remarquables du  $C_{60}$  et des autres fullerènes ont également attiré l'attention de la communauté scientifique. Contrairement au graphite et au diamant, les fullerènes sont solubles dans divers solvants organiques adaptés aux transformations chimiques. Le  $C_{60}$  possède d'intéressantes propriétés photophysiques, notamment la désexcitation quasi-quantitative des états excités singulet en état triplet par croisement inter-système. Les propriétés électrochimiques du  $C_{60}$  sont aussi remarquables. Le  $C_{60}$  peut être facilement réduit (jusqu'à 6 fois par 6 étapes monoélectroniques successives), mais est plus difficilement oxydable.

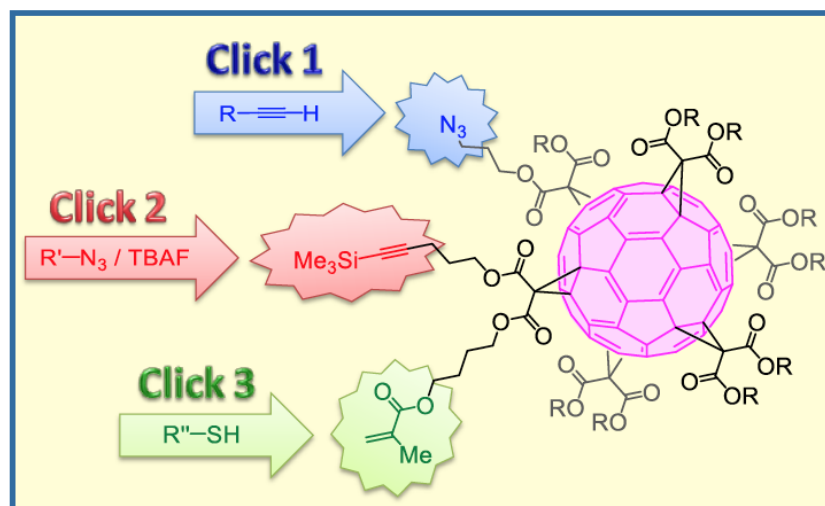
La réactivité du  $C_{60}$  est assimilable à celle d'une polyoléfine électro-déficiente tendue qui peut être le siège de réactions d'additions nucléophiles ou de cycloadditions. L'une des réactions les plus utilisées pour la multi-fonctionnalisation du  $C_{60}$  est la réaction de Bingel.<sup>[3]</sup> Cette réaction consiste à l'addition nucléophile d'un  $\alpha$ -halomalonate sur le  $C_{60}$  et conduit à l'obtention d'un seul dérivé mono-adduit du  $C_{60}$ . Cependant, la poly-fonctionnalisation du  $C_{60}$  est une tâche ardue et est confrontée à des problèmes de régio- et de stéréochimie.<sup>[4,5]</sup> En effet, les dérivés mono-fonctionnalisés de  $C_{60}$  possèdent neuf liaisons 6-6 différentes (liaisons à la jonction entre deux cycles à six atomes) et un mélange de 9 régioisomères est obtenu lors de deux réactions successives sur le cœur fullerénique (Figure 1). La synthèse de tris-adduits du  $C_{60}$  est encore nettement plus difficile car il est théoriquement possible de former jusqu'à 46 régioisomères. L'obtention d'hexa-adduits du  $C_{60}$  d'ordre d'addition octaédrique est fortement conditionnée par la régiochimie des précurseurs bis- et tris-adduits. En effet, seul deux bis-adduits et deux tris-adduits du  $C_{60}$  ont un motif d'addition adéquat pour l'obtention de ces hexa-adduits de  $C_{60}$  (Figure 1).

Les hexa-adduits de fullerène avec une symétrie  $T_h$  et un ordre d'addition octaédrique sont une classe unique de molécules tridimensionnelles.<sup>[6]</sup> Cependant, la synthèse directe d'hexa-adduits de fullerène, à partir de  $C_{60}$  et de malonates, reste difficile et est principalement limité à des malonates relativement simple. Ce problème majeur a été récemment résolu par la synthèse d'hexa-adduits du  $C_{60}$  facilement accessibles, comportant 12 groupes terminaux post-fonctionnalisables, donnant ainsi accès à des systèmes structurellement plus compliqués.<sup>[6-8]</sup> Parmi les réactions de post-fonctionnalisation possibles, la cycloaddition 1,3-dipolaire de Huisgen, entre un azoture et un alcyne vrai, s'est révélé être particulièrement intéressante pour incorporer une large variété de groupes fonctionnels autour du corps fullerénique. Actuellement, le développement d'une

méthodologie efficace pour la synthèse d'hexa-adduits de fullerène comportant deux ou plusieurs fonctions périphériques différentes reste un enjeu majeur.



**Figure 1:** **A gauche:** L'addition d'un second adduit sur un mono-adduit de C<sub>60</sub> peut en principe mener à 9 régioisomères. Relatif au premier adduit, le second peut être localisé dans le même hémisphère (cis), dans l'autre hémisphère (trans) ou à l'équateur (e). **A droite:** Les bis- et tris-adduits du C<sub>60</sub> qui permettent un accès aux hexa-adduits d'ordre d'addition octaédrique.



**Figure 2:** Illustration des possibilités de fonctionnalisation de la sphère du C<sub>60</sub>.

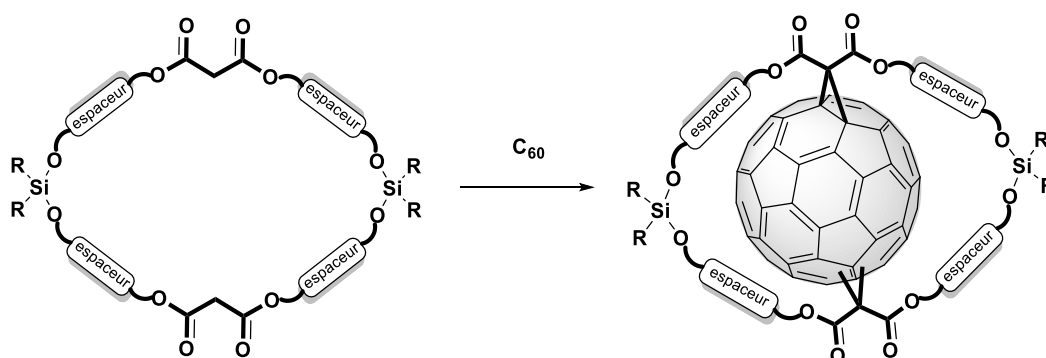
L'objectif de cette thèse sera dans un premier temps de développer des méthodologies de synthèses régiosélectives de bis- et tris-adduits de C<sub>60</sub>. Dans un deuxième temps, il sera question de préparer des synthons C<sub>60</sub> hexa-substitués polyvalents incorporant plusieurs centres réactifs différents et pouvant être sélectivement fonctionnalisés (Figure 2), à partir de

bis- et tris-adduits de  $C_{60}$  adaptés à la formation d'hexa-adduits de  $C_{60}$  d'ordre d'addition octaédrique.

Afin d'obtenir ce type de supports multifonctionnels, il est essentiel de pouvoir accéder facilement à certain des bis- et tris-adduits du  $C_{60}$  possédant un ordre d'addition adéquat ([Figure 1](#)). Dans la littérature, deux approches principales ont été mise au point pour la fonctionnalisation régiosélective du  $C_{60}$ , une approche dite à « bras directeur ou tête directrice » développée par *Diederich et al*<sup>[9]</sup> et une approche macrocyclique développée par *Hirsch et al*.<sup>[10]</sup> Cependant, les travaux effectués jusque-là ne permettaient pas une post-fonctionnalisation aisée de ces dérivés de fullerènes.<sup>[11]</sup> Dans ce contexte, l'utilisation comme tête directrice ou incorporé dans un macrocycle d'une fonction stable et qui peut être facilement déprotégée semble être une idée prometteuse pour une post-fonctionnalisation aisée.

Notre laboratoire s'est donc intéressé à l'utilisation de groupements silylés. Les groupements silylés sont de bons groupes protecteurs de fonctions alcool ou alcyne vrai et peuvent être facilement enlevés par action de fluorure. Les résultats préliminaires obtenus au sein du laboratoire, nous ont conduits à entreprendre une étude plus approfondie du potentiel des groupements silylés.

Dans un premier temps, c'est l'approche macrocyclique qui a été étudiée. La conception du macrocycle est fondée sur trois éléments, (1) des malonates pour l'ancrage sur le  $C_{60}$ , (2) un espaceur pour atteindre une position spécifique sur le  $C_{60}$  et (3) des groupements silylés incorporés dans l'espaceur ([Figure 3](#)).



[Figure 3](#): Approche macrocyclique pour la fonctionnalisation régiosélective du  $C_{60}$ .

Notre étude sur la régiosélectivité des macrocycles bis-malonates s'est axée sur trois paramètres : (1) la taille des macrocycles, (2) la flexibilité/rigidité des espaceurs et (3) le groupe silylé.

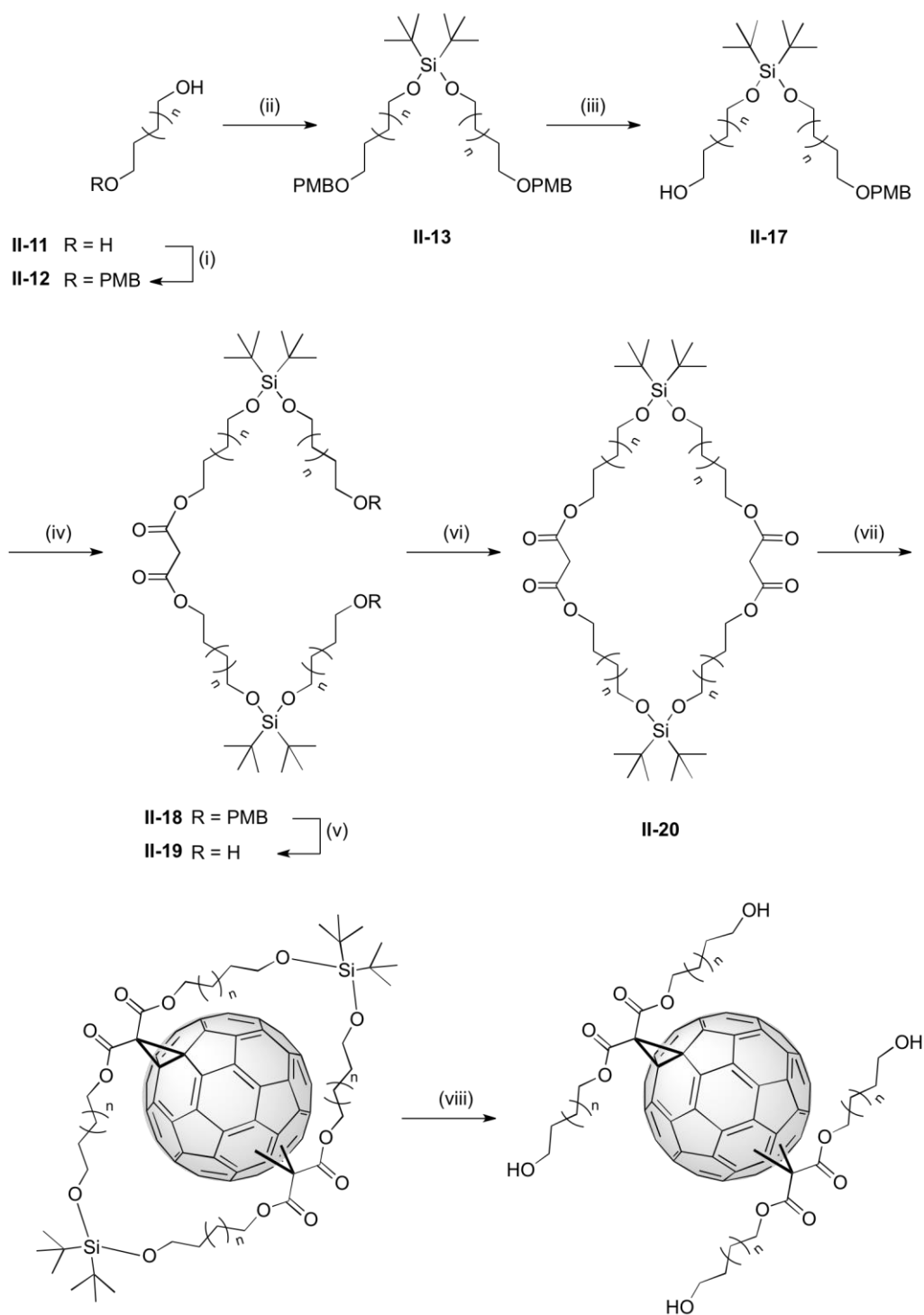
L'élaboration de macrocycles bis-malonates incorporant des groupements silylés, nous a menés vers le développement d'une synthèse dirigée de ces macrocycles à partir d'alcanediols possédant de 4 à 8 atomes de carbone ([Figure 4](#)). Dans un premier temps, les alcanediols sont mono-protégés par des groupes p-méthoxybenzyle. S'en suit une deuxième réaction de protection avec du di-*t*-butylsilyl bis(trifluorométhanesulfonate) pour former une

pince. Cette pince est ensuite mono-déprotégée par réaction avec du 2,3-dichloro-5,6-dicyano-p-benzoquinone (DDQ). Il est à noter que le groupe silylé est inerte dans ces conditions de déprotection. La pince mono-alcool est mise à réagir avec du chlorure de malonyle pour obtenir le mono-malonate, précurseur clé du macrocycle. Une dernière réaction de déprotection avec du DDQ est effectuée pour obtenir le dérivé diol. Finalement, les macrocycles bis-malonates sont obtenus avec de bons rendements (19-50%) par réaction des mono-malonates diols avec du chlorure de malonyle.

Les réactions des macrocycles bis-malonates avec le  $C_{60}$  ont donné de bons résultats et différents régioisomères (*cis-2*, *e* et *trans-3*) ont été obtenus selon la taille des macrocycles. Finalement, une réaction de désilylation par traitement avec un excès de  $BF_3 \cdot Et_2O$  a libéré quatre fonctions alcool pouvant être aisément post-fonctionnalisées.

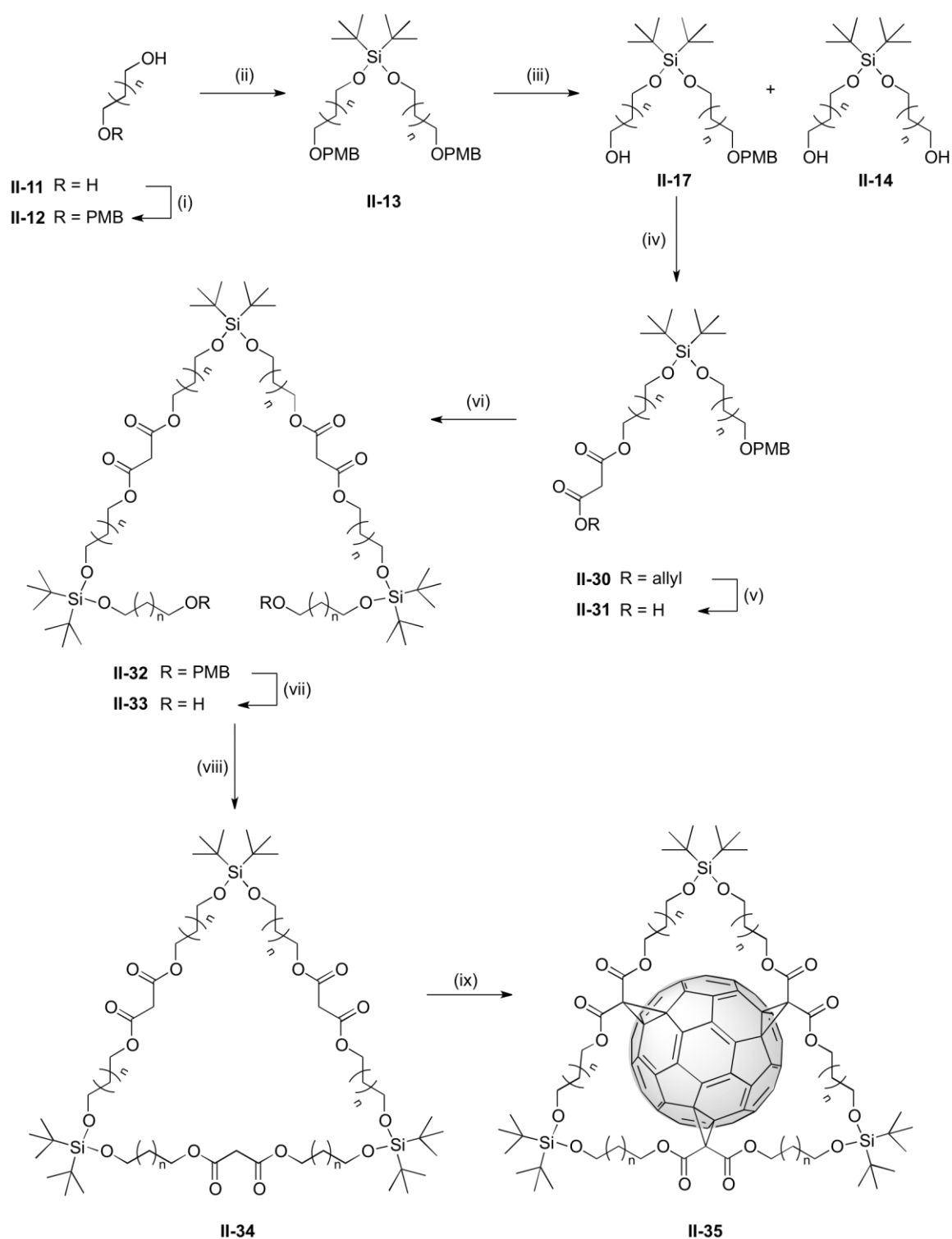
Le contrôle de la régiochimie par notre méthode macrocyclique est à la fois gouverné par la taille du macrocycle et la différence de réactivité des différentes doubles liaisons du  $C_{60}$  après la première fonctionnalisation. Au-delà de la flexibilité/rigidité de l'espaceur, c'est surtout sa longueur et sa forme qui influencent la régiosélectivité mais la fonctionnalisation se fait préférentiellement sur les positions les plus réactives (*e* et *trans-3*). Ce n'est qu'avec les cycles les plus petits que le régioisomère *cis-2* a été obtenue car les positions *e* et *trans-3* sont trop éloignés et ne sont plus accessibles. Le motif O-Si-O joue aussi un rôle non négligeable en apportant une certaine flexibilité au macrocycle. Son hydrolyse permet de générer des synthons tétraalcool. Les bis-adduits *e* formés au cours de cette étude sont de parfaits candidats pour l'élaboration des hexa-adduits de  $C_{60}$  multifonctionnels souhaités.

Notre approche macrocyclique ayant donné de bons résultats pour la formation de bis-adduits du  $C_{60}$ , la synthèse de tris-adduits du  $C_{60}$  a été entreprise en utilisant une stratégie similaire. Pour cela, une synthèse dirigée en 8 étapes de macrocycles tris-malonate a été développée à partir d'alcanediols possédant de 3 à 7 atomes de carbone ([Figure 5](#)). Cette stratégie de synthèse fait intervenir jusqu'à trois groupes protecteurs différents, dont les groupements silylés présent dans la structure finale du macrocycle. Ces trois groupes protecteurs ont dû être précautionneusement choisis afin de pouvoir être sélectivement enlevés en présence des autres groupes protecteurs. Par cette méthode, les réactions de macrocyclisation ont donné de bons rendements (21-63%) pour la formation des macrocycles tris-malonates. Pour deux macrocycles tris-malonates, la réaction avec le  $C_{60}$  s'est avérée être hautement régiosélective et un seul tris-adduit de  $C_{60}$  parmi les 46 régioisomères théoriquement possible a été obtenu. De plus, ces tris-adduits de  $C_{60}$  d'ordre d'addition *e,e,e* ont été obtenus avec d'excellents rendements (39 et 61%).



**Figure 4: Réactifs et conditions** :  $n = 1-5$  ; (i) PMBCl, Ag<sub>2</sub>O, CH<sub>2</sub>Cl<sub>2</sub>, t.a., 48 h (69-95%); (ii) tBu<sub>2</sub>Si(OTf)<sub>2</sub>, DMF, imidazole, t.a., 12 h (60-91%); (iii) DDQ (1 éq), CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, t.a., 6 h (47-50%); (iv) chlorure de malonyle, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, t.a., 2 h (67-96%); (v) DDQ (2.5 éq), CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, t.a., 2 h (68-85%); (vi) chlorure de malonyle, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, t.a., 2 h (19-50%); (vii) C<sub>60</sub>, I<sub>2</sub> and DBU, PhMe, rt, 1 h; (viii) BF<sub>3</sub>.Et<sub>2</sub>O, CH<sub>2</sub>Cl<sub>2</sub>, CH<sub>3</sub>CN, rt, 12 h.





**Figure 5: Réactifs et conditions :** (i) PMBCl, Ag<sub>2</sub>O, CH<sub>2</sub>Cl<sub>2</sub>, t.a., 12 h; (ii) tBu<sub>2</sub>Si(OTf)<sub>2</sub>, DMF, pyridine, t.a., 12 h (iii) DDQ (1 éq), CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, t.a., 1 h; (iv) **II-28**, DCC, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, 0°C to t.a., 12 h; (v) Pd(PPh<sub>3</sub>)<sub>4</sub>, morpholine, THF, t.a., 4 h; (vi) **II-14b-e**, DCC, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, 0°C à t.a., 12 h; (vii) DDQ, CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, t.a., 6 h; (viii) chlorure de malonyle, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, t.a., 24 h; (ix) C<sub>60</sub>, I<sub>2</sub>, DBU, PhMe, t.a., 1 h.

L'approche macrocyclique s'est avérée être très intéressante pour la fonctionnalisation régiosélective du  $C_{60}$  et les produits d'addition sont généralement obtenus avec de bons rendements. Cependant, dans le cas des tris-malonates macrocycliques, le grand nombre d'étapes pour la synthèse des macrocycles ne donne pas un accès rapide au tris-adduits *e,e,e*.

Ceci nous a conduits à développer une approche alternative de type « tête directrice ». Celle-ci repose sur la réaction du  $C_{60}$  avec un tris-malonate construit autour d'un motif trialcoxysilane.

Dans ce cas, la préparation des précurseurs tris-malonate est aisée. Elle est réalisée dans un premier temps (Figure 6), par une réaction d'estérification entre du chlorure d'éthyle malonate et un diol. Les mono-malonate-alcools ainsi obtenus sont traités avec du *t*-BuSiCl<sub>3</sub> pour former les tripodes tris-malonate correspondant. La faible stabilité de ces produits, nous a amené à effectuer la triple réaction de Bingel avec le  $C_{60}$  dans des conditions modifiées. En particulier, il est important de maintenir la température du milieu réactionnel en dessous de -5°C pour éviter des réactions secondaires impliquant le groupement trialcoxysilane. Pour 5 des 8 espaceurs testés, le tris-adduit *e,e,e* a pu être facilement isolé sous forme d'un mélange racémique. La déprotection du groupe silylé a ensuite été effectuée afin d'obtenir des tris-adduits triols post-fonctionnalisables (Figure 6).

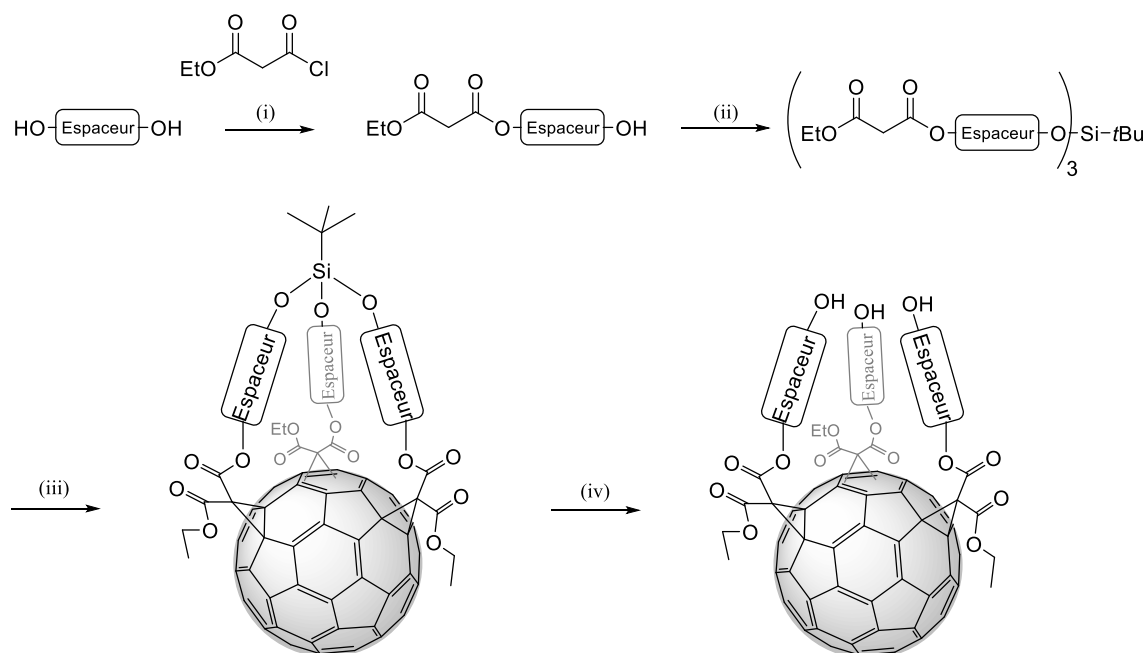
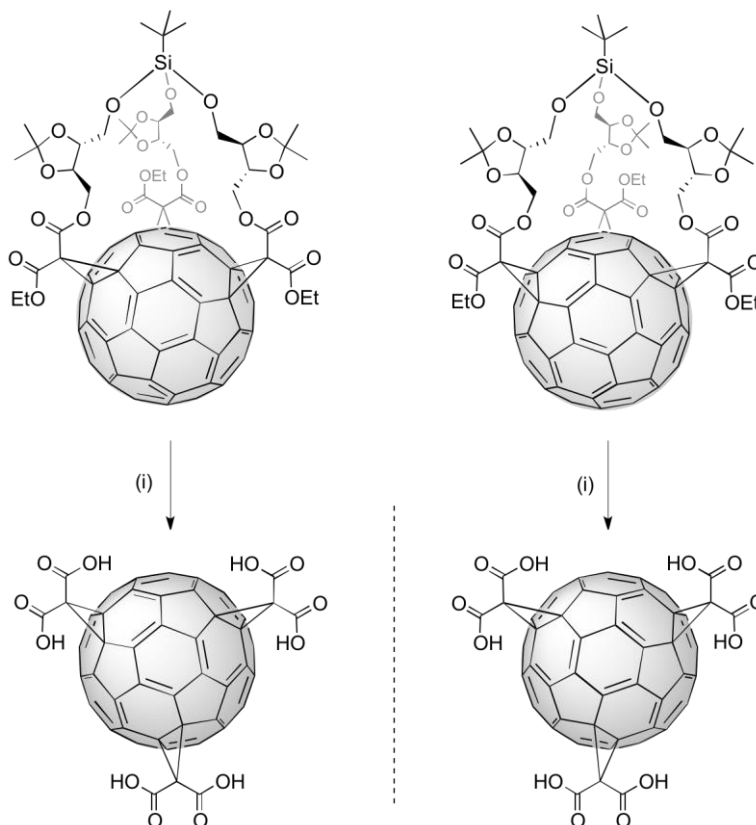


Figure 6 : **Réactifs et conditions** : (i) pyridine, THF, 0°C à t.a., 12 h; (ii) *t*BuSiCl<sub>3</sub>, imidazole, DMF, t.a., 12 h; (iii)  $C_{60}$ , I<sub>2</sub>, DBU, toluène, -15°C, 1 h.

Cette méthodologie de synthèse hautement régiosélective a aussi été utilisée pour la préparation de tris-adduits *e,e,e* optiquement pur. Pour cela des diols optiquement purs ont été utilisés selon les mêmes conditions de réaction. La différence par rapport à la synthèse des tris-adduits présentée précédemment est que l'on forme un mélange de

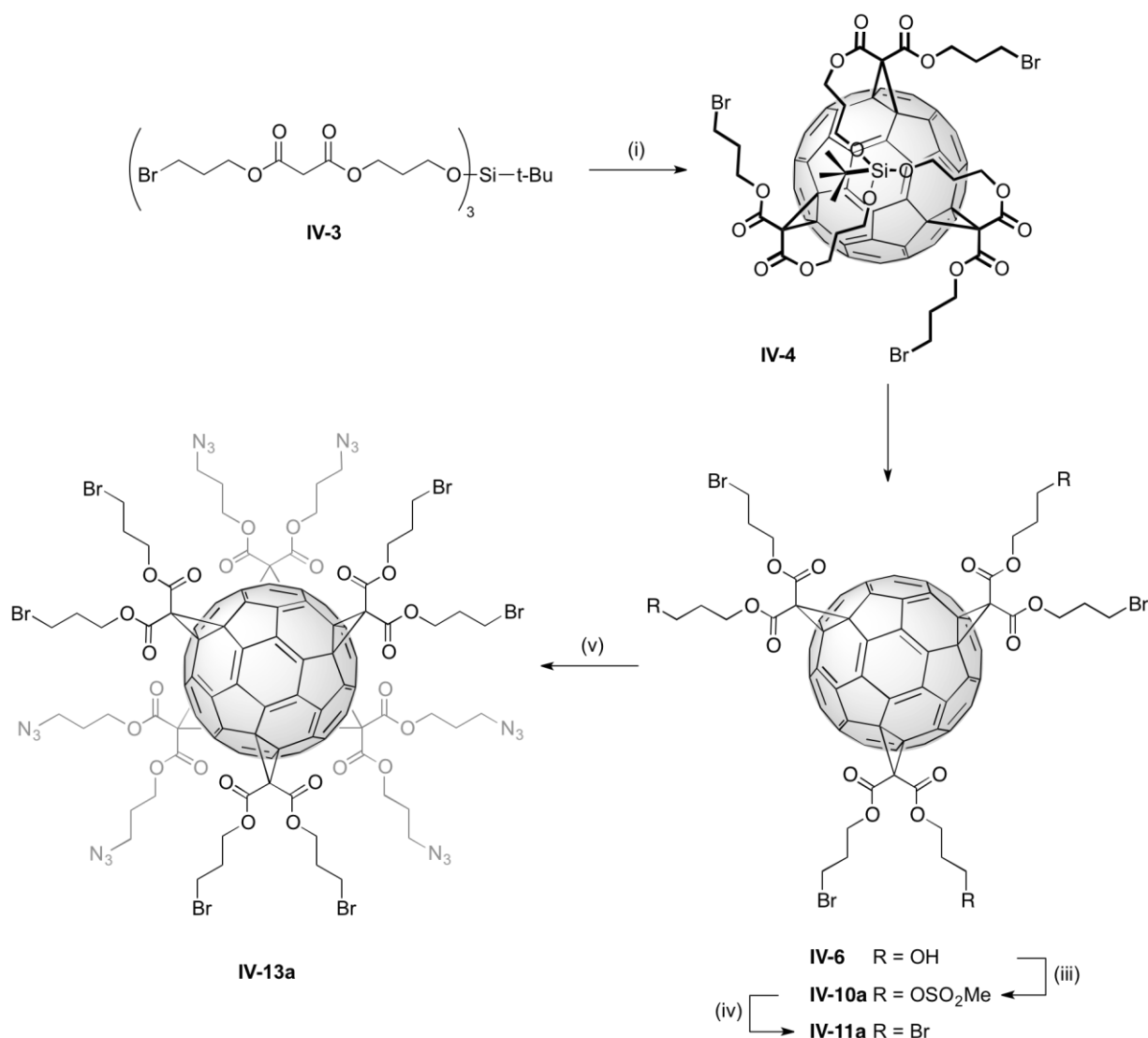
diastéréoisomères et non plus un mélange d'énantiomères. La différence de polarité entre ces diastéréoisomères permet leur séparation par une chromatographie sur colonne de silice classique. L'hydrolyse des esters maloniques a finalement conduit à l'obtention de tris-adduits *e,e,e* optiquement purs et comportant 6 fonctions acides carboxyliques (Figure 7).



**Figure 7: Réactifs et conditions :** (i) PhMe, MeOH, NaH, 60 °C.

Cette stratégie « tête directrice » fournit un accès rapide, simple, régio- et stéréosélectif à des tris-adduits *e,e,e* approprié à la formation d'hexa-adduits d'ordre d'addition octaédrique. La synthèse des synthons hexa-adduits multifonctionnels a dès lors pu être entreprise.

Dans le but d'avoir des synthons  $C_{60}$  hexa-adduits les plus fonctionnalisables possibles, une modification de la chaîne latérale des malonates a été effectuée dans un premier temps (Figure 8). La transformation des synthons bis- et tris-adduits a ensuite été effectuée de manière à permettre deux réactions « click » successives après la formation de l'hexa-adduits de  $C_{60}$ . Au total, trois synthons « cliquable » d'hexa-adduits mixtes de  $C_{60}$  ont été préparé. Par deux réactions « click » successives, l'introduction de deux types de fonctions différentes à la périphérie des hexa-adduits de  $C_{60}$  a été effectuée avec de très bons rendements (75-99%). Ces hexa-adduits de  $C_{60}$  proposent différents ratio de fonction « cliqué » (3:6 ; 6:6 et 4:8) dans deux zones distinctement prédéfini.



**Figure 8: Réactifs et conditions :** (i) C<sub>60</sub>, I<sub>2</sub>, DBU, PhMe, -15 °C, 1 h (13%); (ii) BF<sub>3</sub>.Et<sub>2</sub>O, CH<sub>2</sub>Cl<sub>2</sub>/MeCN 2:1, t.a., 12 h (86%). (iii) MeSO<sub>2</sub>Cl, Et<sub>3</sub>N, CH<sub>2</sub>Cl<sub>2</sub>, 0 °C, 2 h (86%); (iv) LiBr, THF, 60 °C, 12 h (83%); (v) Malonate azoture, CBr<sub>4</sub>, DBU, PhMe, t.a., 12 h (42%).

La post-fonctionnalisation des fonctions alcools du synthon tris-adduit triol (**IV-6**) peut aussi être effectuée par une réaction d'estérification. Cela permet une différenciation supplémentaire des types de fonctions à la périphérie des hexa-adduits (Figure 9). C'est ainsi qu'un troisième hexa-adduits [3:3]-(3:9) à deux types de fonctions a pu être obtenu. Cette méthodologie a ensuite été étendue à la synthèse d'hexa-adduits à trois types de fonctions. Par une réaction d'estérification suivie de deux réactions « click », la post-fonctionnalisation d'un quatrième hexa-adduit [3:3]-(3:3:6) a été achevée. La formation de trois hexa-adduits [3:3] structurellement différent montre la polyvalence de notre méthodologie. Il est aussi à noter que tous les hexa-adduits présentés dans ce travail ont été synthétisé avec un contrôle de la régiochimie et de la stéréochimie, définissant des zones distinctes de fonctionnalités (Figure 9).

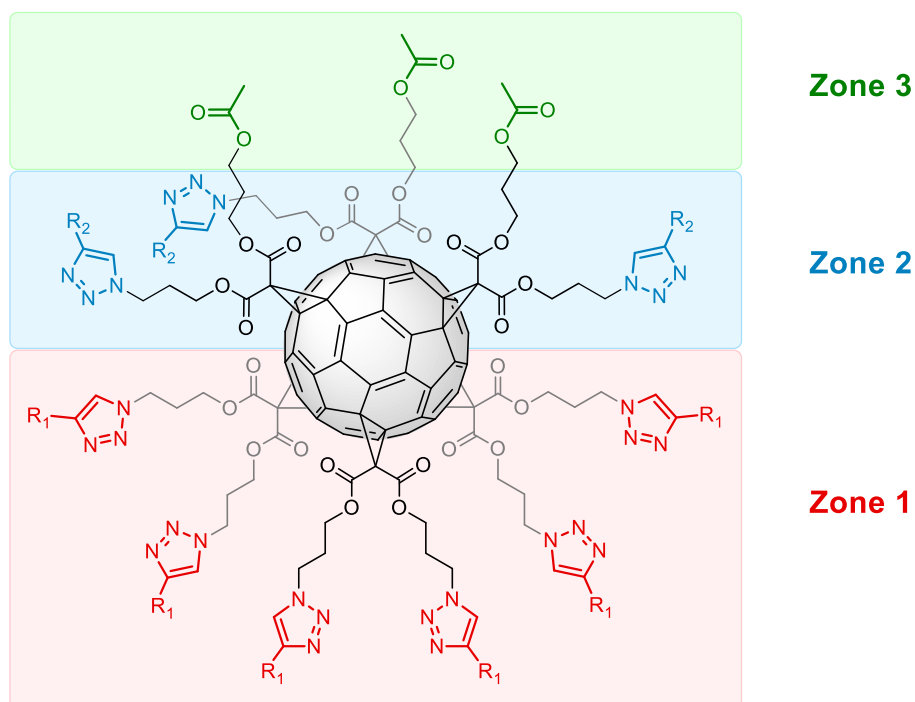
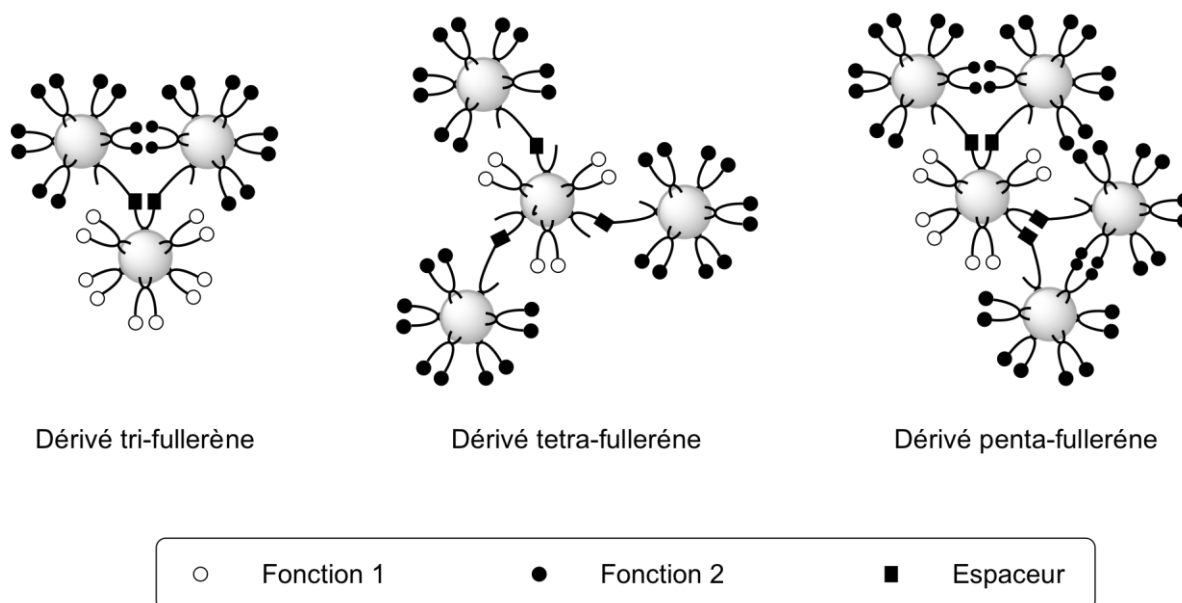


Figure 9: Illustration des trois de distinctes de fonctionnalités de l'hexa-adduit [3:3]-(3:3:6).

A partir des synthons d'hexa-adduits mixtes de  $C_{60}$  développés précédemment, l'élaboration d'édifices hyperfonctionnels comportant plusieurs hexa-adduits de  $C_{60}$  autour d'un hexa-adduit de  $C_{60}$  central a été effectuée.

Dans un premier temps c'est les synthèses de dérivés tri-, tétra- et penta-fullerènes qui ont été réalisées (Figure 10). Pour des raisons d'accessibilité à ces systèmes hyperfonctionnels, un espaceur comportant deux fonctions alcynes vrai à ses extrémités a dû être préparé, afin de pouvoir relier les hexa-adduits périphériques à l'hexa-adduit central. L'utilisation de la cycloaddition 1,3-dipolaire de type Huisgen a permis l'obtention de ces dérivés tri-, tétra- et penta-fullerènes avec de bon rendements. Il est à noter que ces dérivés comportent deux sortes de fonctions, celles de l'hexa-adduit central et celles des hexa-adduits périphériques.

Dans la continuité de ces résultats, un « mégaball » mixte a aussi été préparé (Figure 11). Ce dernier a été préparé à partir d'un hexa-adduit du  $C_{60}$  de type [2:4] comme cœur dendritique. Cet hexa-adduit central est fonctionnalisé par deux réactions successives de cycloaddition 1,3-dipolaire de type Huisgen avec des hexa-adduits de  $C_{60}$  fonctionnalisés par deux types de fonctions. La première réaction « click » introduit 80 unités de la fonction 1 répartis uniformément sur 8 hexa-adduits périphériques. La deuxième réaction « click » introduit quant à elle 40 unités de la fonction 2. Le mégaball mixte ainsi formé comporte au total 120 fonctions de deux types à sa périphérie.



*Figure 10: Edifices hyperfonctionnels préparés par assemblage d'hexa-adduits de C<sub>60</sub>.*

## **Conclusion.**

Dans un premier temps, la bis- et tris-fonctionnalisation du C<sub>60</sub> a été effectuée par une approche macrocyclique. Différents bis- et tris-adduits du C<sub>60</sub> ont été obtenus avec de bonnes régiosélectivités. La régiochimie de notre méthodologie est gouvernée par la taille des macrocycles et par la présence de groupes silylés au sein de la structure des macrocycles. Les dérivés polyols obtenus par déprotection des groupes silylés ouvrent de nouvelles perspectives pour la post-fonctionnalisation de multi-adduits de C<sub>60</sub>.

Dans un deuxième temps, la tris-fonctionnalisation du C<sub>60</sub> a été effectuée par une approche « tête directrice ». Par cette approche, un accès rapide, simple, régio- et stéréosélectif à des tris-adduits *e,e,e* approprié à la formation d'hexa-adduits d'ordre d'addition octaédrique a été obtenu. La synthèse et la séparation de tris-adduits *e,e,e* optiquement purs ont également été réalisées.

A partir de nos méthodologies de synthèse de bis- et tris-adduits de C<sub>60</sub>, trois synthons d'hexa-adduits mixtes de C<sub>60</sub> ont été préparés. La post-fonctionnalisation de ces synthons a été effectuée par réaction de cycloaddition 1,3-dipolaire de type *Huisgen* mais aussi par réaction d'estérification, afin d'obtenir des hexa-adduits de C<sub>60</sub> multifonctionnels avec de bons rendements. Le contrôle de la régio- et stéréochimie de notre méthodologie permet de définir jusqu'à trois zones distinctes comportant chacune des fonctions différentes. La post-fonctionnalisation des synthons par chimie « click » donne accès à la préparation contrôlée de nouveaux nanomatériaux globulaires multifonctionnels.

Dans la dernière partie de ce travail de thèse, la méthodologie de synthèse d'hexa-adduits de C<sub>60</sub> multifonctionnels a été mise à profit pour l'élaboration d'édifices hyperfonctionnels comportant plusieurs hexa-adduits de C<sub>60</sub> autour d'un hexa-adduit de C<sub>60</sub>

central. Cette approche dendritique est une voie d'accès à l'élaboration contrôlée de nano-édifices multifonctionnels.

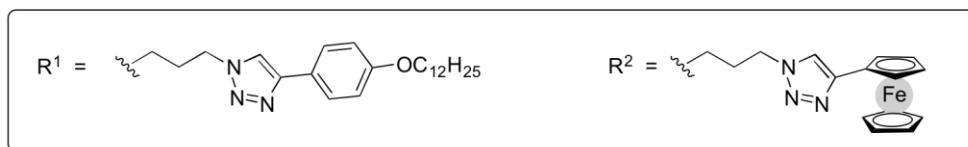
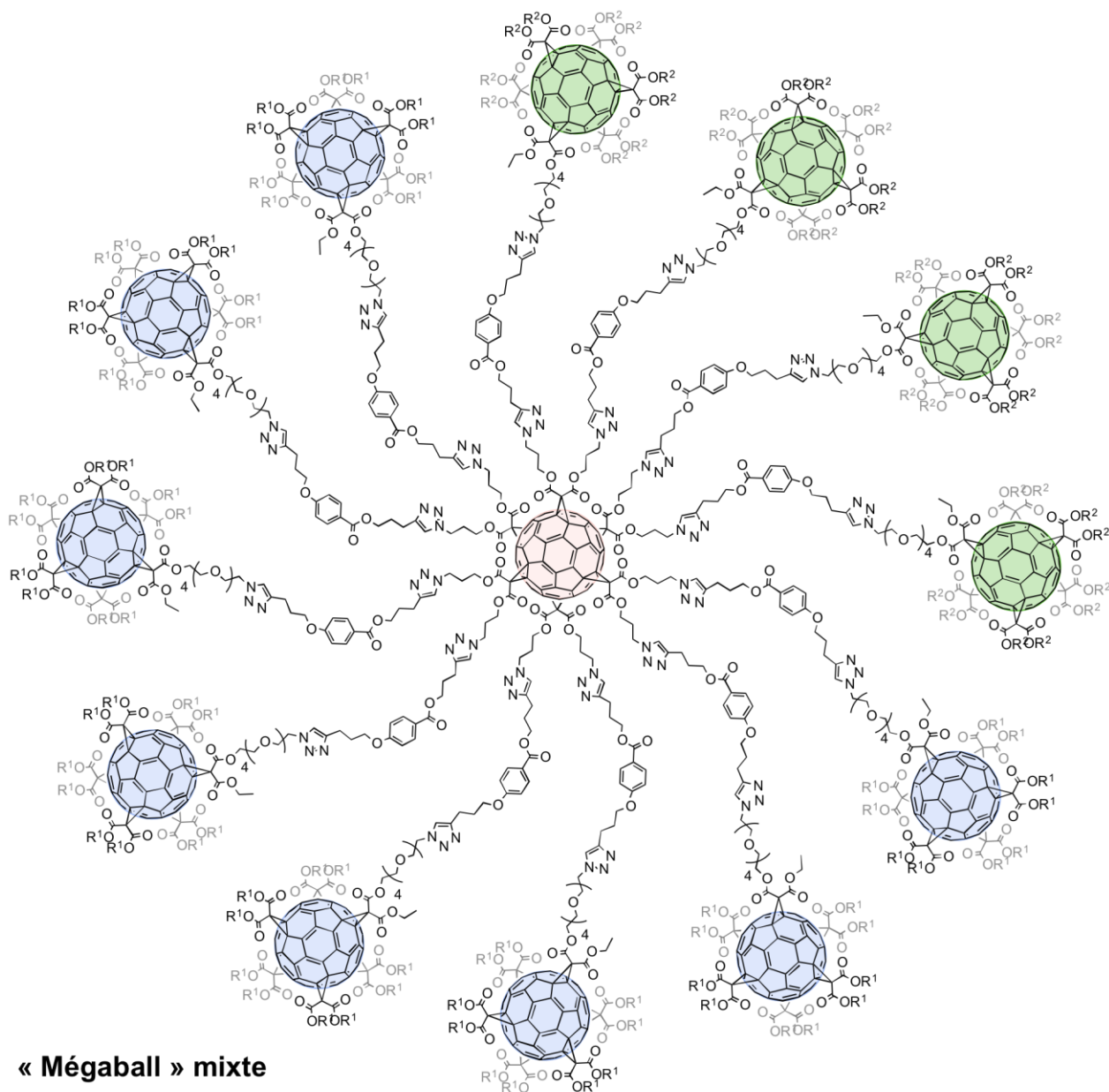


Figure 11: Structure du mégaball mixte comportant au total 120 fonctions de deux types à sa périphérie.

Références :

- [1] H. W. Kroto, J. R. Heath, S. C. O'Brien, R. F. Curl, R. E. Smalley, *Nature* **1985**, *318*, 162–163.
- [2] W. Krätschmer, L. D. Lamb, K. Fostiropoulos, D. R. Huffman, *Nature* **1990**, *347*, 354–358.
- [3] C. Bingel, *Chem. Ber.* **1993**, *126*, 1957–1959.
- [4] A. Hirsch, in *Fuller. Relat. Struct.* (Ed.: P.D.A. Hirsch), Springer Berlin Heidelberg, **1999**, pp. 1–65.
- [5] C. Thilgen, F. Diederich, *Chem. Rev.* **2006**, *106*, 5049–5135.
- [6] J. Iehl, R. P. de Freitas, B. Delavaux-Nicot, J.-F. Nierengarten, *Chem. Commun.* **2008**, 2450–2452.
- [7] J. Iehl, J.-F. Nierengarten, *Chem. – Eur. J.* **2009**, *15*, 7306–7309.
- [8] I. Nierengarten, J.-F. Nierengarten, *Chem. Rec.* **2015**, *15*, 31–51.
- [9] L. Isaacs, R. F. Haldimann, F. Diederich, *Angew. Chem. Int. Ed. Engl.* **1994**, *33*, 2339–2342.
- [10] U. Reuther, T. Brandmüller, W. Donaubauer, F. Hampel, A. Hirsch, *Chem. – Eur. J.* **2002**, *8*, 2261–2273.
- [11] C. Thilgen, F. Diederich, *Comptes Rendus Chim.* **2006**, *9*, 868–880.



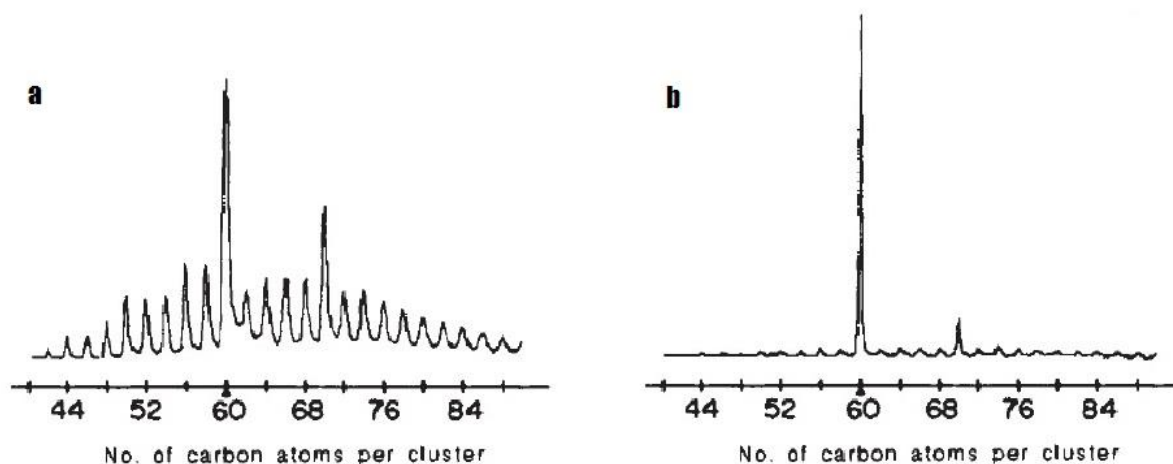


# **Chapter I : General introduction.**

# 1. Generalities.

## 1.1 Discovery of a new allotropic form of carbon.

Fullerenes are caged molecules exclusively constituted by carbon. After diamond and graphite, they represent the third allotropic form of carbon. The history of this family of compounds began in 1966 when D. E. H. Jones proposed for the first time the idea of large hollow carbon cages<sup>[1]</sup>, later known as giant fullerenes. In the same year, a premise of the structure of fullerenes was drawn by the synthesis of corannulene by G. Lawton et al.<sup>[2]</sup> Stimulated by the discovery of corannulene, Osawa proposed in 1970 a truncated icosahedral structure for an hypothetical  $[C_{60-I_h}]$  fullerene but this structure remained theoretical for many years.<sup>[3]</sup> The first carbon clusters were produced in 1984 by laser vaporization of graphite. Time-of-flight mass spectrometry analysis revealed that only carbon clusters  $C_n$  with an even number and  $n = 40-190$  could be observed.<sup>[4]</sup> However no specific structure could be proposed for the observed carbon clusters despite the fact that  $C_{60}$  was among them. The breakthrough occurred one year later (1985) when Kroto, Curl and Smalley published a Nature article<sup>[5]</sup> in which they described the production of fullerenes using the same technic.<sup>[4]</sup>  $C_{60}$  and  $C_{70}$  were the most abundant fullerenes forms (Figure I-1-a). By optimizing the experimental conditions, they were able to predominantly produce clusters constituted by 60 C atoms (Figure I-1-b). They rationalized these results to a better stability of the  $C_{60}$  cluster and proposed a truncated icosahedral structure to explain this extra-stability.

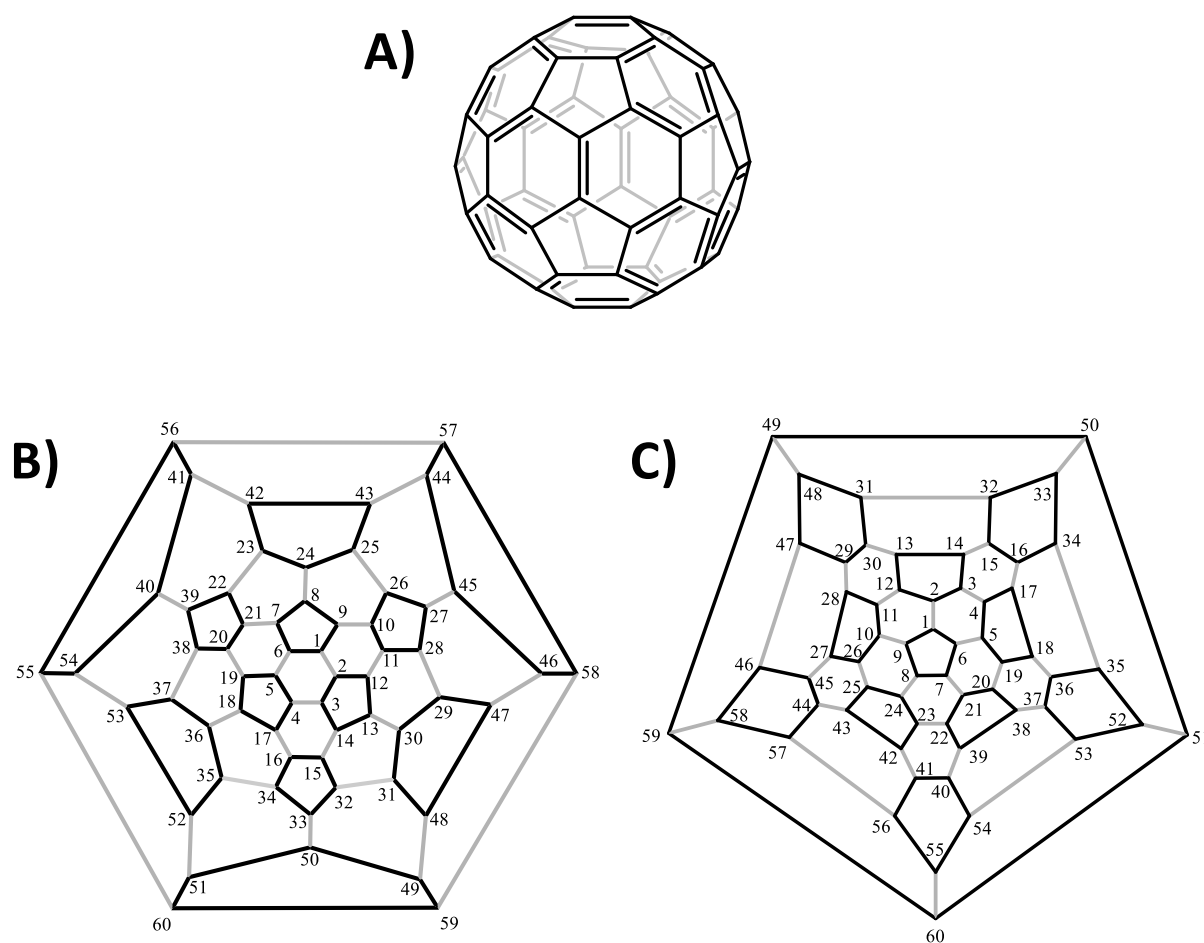


*Figure I-1: "Time of flight mass spectra of carbon clusters prepared by laser vaporization of graphite and cooled in a supersonic beam. a) The spectrum was obtained when roughly 760 torr helium was present over the graphite target at the time of laser vaporization. b) The spectrum was obtained by maximizing these cluster thermalization and cluster-cluster reactions."<sup>[5]</sup>*

A new allotropic form of carbon was born. The most abundant fullerene is  $C_{60}$  also called buckminsterfullerene in reference of the geodesic domes created by the architect Buckminster Fuller. After the discovery of other clusters (e.g.  $C_{70}$ ,  $C_{76}$ ,  $C_{80}$  and  $C_{82}$ ) the shortest name "Fullerene" was kept to describe this family of compounds.

Two important points contributed to the development of fullerene chemistry: (i) unlike graphite or diamond that are polymeric and insoluble, fullerenes are well-defined molecules soluble in various organic solvents and (ii) their gram-scale production developed by Krätschmer, Huffman et al. in 1990 made them available to the scientific community.<sup>[6]</sup>

With the development of fullerene chemistry a unified nomenclature was needed. The International Union of Pure and Applied Chemistry (IUPAC) recommended in 2002 the use of  $(C_{60-h})[5,6]$  fullerene as systematic name. The number of carbon atoms and the symmetry group of the cluster and the ring size (square bracketed number) in the fullerene are indicated.<sup>[7]</sup> The recommended systematic numbering of  $C_{60}$  is shown in [Figure I-2](#).

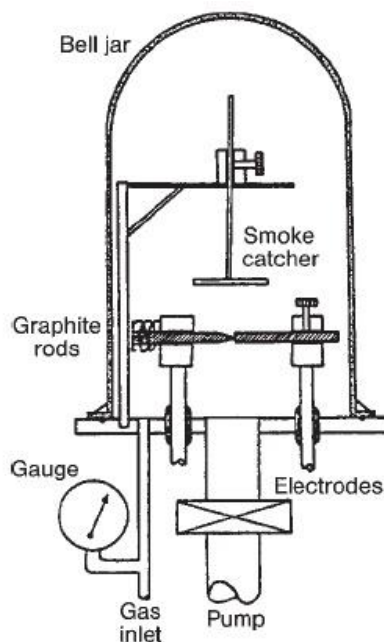


*Figure I-2 : Schematic representations of  $C_{60}$ . A) Lewis structure; B) Schlegel diagram (hexagon centered) with systematic numbering. ; C) Schlegel diagram (pentagon centered) with systematic numbering.*

## 1.2 Production and Purification.

The macroscopic production of  $C_{60}$  was first achieved in 1990 by Krätschmer et al by resistive heating of graphite.<sup>[6]</sup> In a bell jar under an atmosphere of  $\sim 133$  mbar of helium, they connected two graphite rods in close contact to two copper electrodes ([Figure I-3](#)). For

producing soot, an electric current was applied to evaporate the graphite rods by resistive heating. Purification of the resulting soot afforded a fullerene yield of 5-10%.



*Figure I-3 : Fullerenes production system used by Krätschmer et. al. (from "Fullerenes Chemistry and Reaction", Andreas Hirsch and Michael Brettreich, Wiley-VCH, 2002).*

Some other techniques more or less efficient were also developed to produce fullerenes such as arc heating of graphite<sup>[8]</sup>, solar generators<sup>[9]</sup> or inductive heating of graphite.<sup>[10]</sup>

Currently, the most important technique to produce fullerenes is by combustion in optimized sooting flames.<sup>[11-13]</sup> Fullerenes in hydrocarbon flames were first revealed in 1987 by mass spectrometry.<sup>[14]</sup> Continuous optimization of this technique allowed later industrial production of fullerenes.

Isolation of fullerenes from soot can be achieved by extraction<sup>[6]</sup> or by sublimation<sup>[15]</sup>. Originally purified from soot by extraction in benzene, only a mixture of fullerenes was recovered.<sup>[6]</sup> Several methods were then developed to separate fullerenes implying especially chromatography techniques.<sup>[16]</sup> A common method is the use of chromatography on alumina, but it can require a big amount of alumina and solvent.<sup>[17,18]</sup> The use of Soxhlet-chromatography has considerably improved the efficiency of the separation on alumina.<sup>[19,20]</sup> Another efficient and inexpensive method is chromatography on mixtures of charcoal and silica gel.<sup>[21]</sup> Instead of chromatography, fractional crystallization can also be used to purify C<sub>60</sub>.<sup>[22]</sup>

The development of production and purification methods has allowed industrial scale production of fullerenes. Companies such as *Frontier Carbon Corporation* have produced several tons of fullerenes using an industrial process based on combustion in sooting flames. Concurrently, the cost of fullerenes has considerably decreased from about 5000 \$/g at the origin to an affordable price of 20-25 \$/g (for C<sub>60</sub>) nowadays.

## 1.3 Fullerene C<sub>60</sub>: Structure and Properties.

### 1.3.1 Structure

C<sub>60</sub> Buckminsterfullerene is the smallest stable fullerene and the most abundant. As all fullerenes, it has a ball shape structure and is formed by 20 hexagons and 12 pentagons. Even before its discovery, a truncated icosahedral structure was proposed<sup>[3]</sup>. Its experimental discovery<sup>[5]</sup> highlighted the “extra stability” of C<sub>60</sub>. It has been demonstrated that the more favorable and stable structure is an arrangement with all twelve pentagons isolated by hexagons, leading to the “Isolated Pentagon Rule” (IPR)<sup>[23,24]</sup>. The juxtaposition of two pentagons increases the strain energy and destabilizes the structure. On the other hand, more the structure is symmetrical and more the strain is uniformly distributed. The structure of choice was therefore a truncated icosahedron. Theoretical calculations were in agreement with this structure<sup>[25,26]</sup> and it was experimentally confirmed by several technics<sup>[27–30]</sup>. In particular, the <sup>13</sup>C NMR spectrum of C<sub>60</sub> showing only one signal for the 60 equivalent carbon atoms was in agreement with its high symmetry (*I<sub>h</sub>*).

The diameter of [C<sub>60</sub>-*I<sub>h</sub>*]fullerene is 10.34 Å if the π electron cloud is considered. At room temperature the C<sub>60</sub> molecules adopt a face-centered cubic (fcc) organization<sup>[28]</sup>. The spherical shape of C<sub>60</sub> implies a pyramidalization of the sp<sup>2</sup> carbons. A consequence of this geometry is a weak π orbital overlap and no real delocalization of the π-electrons preventing a “superaromaticity”. Two different bonds can be distinguished (Figure I-4). A [5,6] bond of 1.45 Å at the junction of a pentagon and a hexagon, and a [6,6] bond of 1.38 Å at the junction of two hexagons. The [5,6] bond has a simple bond character and the [6,6] bond has a double bond character. The reactivity of the molecule is centered on this last bond.

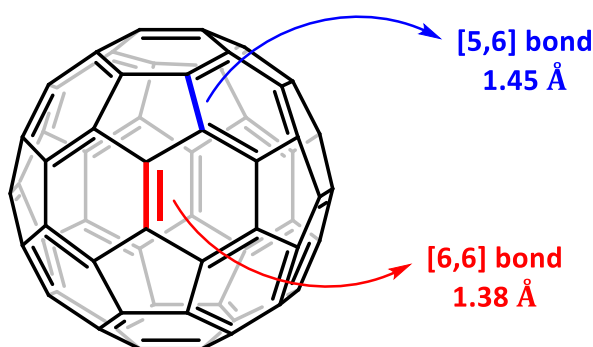


Figure I-4 : Representation of C<sub>60</sub> and its two different type of bonds.

### 1.3.2 Solubility

Solubility is an important parameter to take into account for the chemical transformation of compounds. Systematic studies have highlighted the solubility of C<sub>60</sub> in a large variety of solvents (Table I-1)<sup>[31]</sup>. The aromatic solvents are the best to solubilize C<sub>60</sub>, especially naphthalene derivatives. The most used solvents for fullerene chemistry are 1,2-dichlorobenzene and toluene with a solubility of 27 and 2.8 mg/mL respectively. C<sub>60</sub> is not

soluble in polar solvents and only slightly soluble in alkanes or haloalkanes. The low solubility of  $C_{60}$  is related to strong  $\pi$ - $\pi$  interactions and its tendency to aggregate. Even in the best solvent, 1-chloronaphthalene,  $C_{60}$  is only moderately soluble (51 mg/mL).

*Table I-1 : Solubility of  $C_{60}$  in various solvents*

Solvent	[ $C_{60}$ ], mg/mL	mole fraction ( $\times 10^4$ )
<b>alkanes</b>		
n-pentane	0,005	0,008
n-hexane	0,043	0,073
cyclohexane	0,036	0,059
n-decane	0,071	0,19
decalins	4,6	9,8
<b>haloalkanes</b>		
dichloromethane	0,26	0,27
chloroform	0,16	0,22
carbon tetrachloride	0,32	0,4
1,1,2,2-tetrachloroethane	5,3	7,7
<b>polars</b>		
methanol	0,000	0,000
ethanol	0,001	0,001
acetone	0,001	0,001
acetonitrile	0,000	0,000
<b>benzenes</b>		
benzene	1,7	2,1
toluene	2,8	4
benzonitrile	0,41	0,71
chlorobenzene	7	9,9
1,2-dichlorobenzene	27	53
1,2,4-trichlorobenzene	8,5	15
<b>naphthalenes</b>		
1-methylnaphthalene	33	68
1-phenylnaphthalene	50	131
1-chloronaphthalene	51	97
<b>miscellaneous</b>		
carbon disulfide	7,9	6,6
tetrahydrofuran	0,000	0,000
pyridine	0,89	0,99

### 1.3.3 Photophysical properties.

The photophysical studies of  $C_{60}$  in solution have revealed some of its remarkable properties that can most certainly be attributed to its unique geometry<sup>[32]</sup>. Only a very weak fluorescence was observed for  $C_{60}$ <sup>[33]</sup>. This is due to a nearly quantitative InterSystem Crossing (ISC) from the Singlet excited state to the Triplet excited state. Despite a very high Triplet

quantum yield ( $\Phi_T = 0.96$ ), no phosphorescence could be observed. The deactivation of the Triplet excited states occurs *via* non-radiative deactivation and/or bimolecular quenching depending on the experimental conditions. In the presence of molecular oxygen, very efficient quenching by  $O_2$  leading to  $^1O_2^*$  occurs. On the other hand, electron transfer is observed in the presence of donor molecules. These deactivation processes are important for various applications in material science<sup>[34,35]</sup> or in biology<sup>[36]</sup>. The absorption spectrum of  $C_{60}$  is characterized by a strong absorption in UV with representative bands at 211 nm, 256 nm and 328 nm (Figure I-5). The purple color of  $C_{60}$  is due to forbidden transitions between 410 and 620 nm<sup>[17]</sup>.

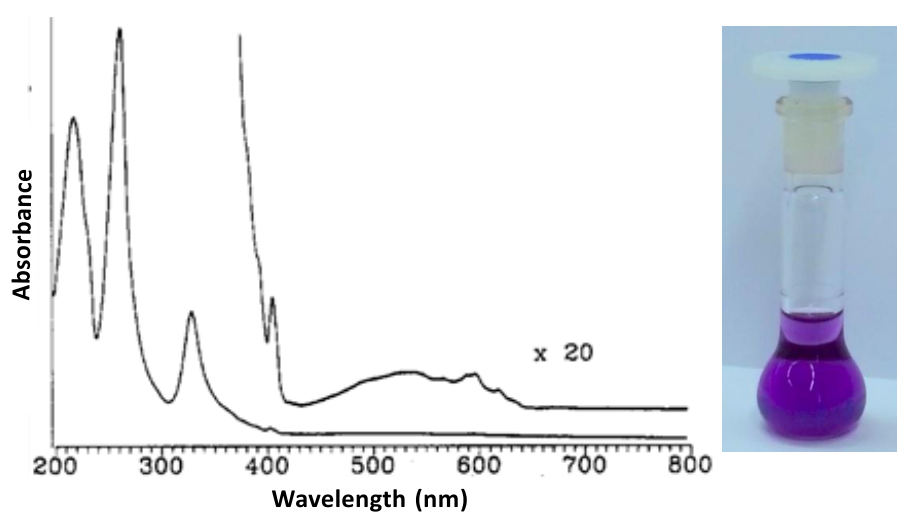


Figure I-5 : Absorption spectrum of  $C_{60}$  in *n*-hexane.

#### 1.3.4 Electrochemical properties.

Huckel molecular orbital (HMO) calculations shows a triply degenerated LUMO orbitals of low energy.<sup>[37]</sup> In theory  $C_{60}$  should be able to accept up to 6 electrons. Six mono-electronic reversible reduction of  $C_{60}$  were effectively reported in 1992 by Echegoyen et al.<sup>[38]</sup> The potentials measured for the six electron reductions relative to  $Fc/Fc^+$  were -0.98, -1.37, -1.87, -2.35, -2.85, -3.26 V in  $CH_3CN$ /toluene at -10 °C.<sup>[38]</sup> According to the authors,  $C_{60}^{5-}$  appears to be stable in solution at room temperature and  $C_{60}^{6-}$  at -10 °C.

If the reduction of  $C_{60}$  is relatively easy, its oxidation is more difficult and requires strict conditions. Oxidation of  $C_{60}$  up to  $C_{60}^{3+}$  was reported in 2003 by Bruno et al.<sup>[39]</sup> To be able to generate  $C_{60}^{3+}$  species they performed CV experiments at -55 °C with ultra-dry  $CH_2Cl_2$  and TBAAsF<sub>6</sub> as electrolytes which has a high oxidation resistance and low nucleophilicity.  $C_{60}^{2+}$  and  $C_{60}^{3+}$  are unstable and react easily to form electroactive materials.



## 2. Reactivity of C<sub>60</sub> and its derivatives.

### 2.1 Introduction.

The unique geometry of [C<sub>60</sub>-I<sub>h</sub>]Buckminsterfullerene is at the basis of its remarkable reactivity made from strain and electronic arguments. The spherical shape of C<sub>60</sub> implies a pyramidalization of the carbon atoms. Deviation from planarity modifies the hybridization of the sp<sup>2</sup> σ and π orbitals. It was calculated by π orbital axis vector (POAV) analysis that σ orbitals have a sp<sup>2.278</sup> hybridization and π orbitals a s<sup>0.09</sup>p hybridization approximately.<sup>[37]</sup> This rehybridization disables the aromaticity due to unsuitable orbital geometry and electronic effects. In addition, it also impacts π\* orbitals which are lower in energy and exhibit a “considerable s character”.<sup>[40]</sup> With a low lying LUMO level, C<sub>60</sub> can be easily reduced and be considered as an electronegative molecule. Its electron acceptor character makes it suitable for addition and redox reactions. Moreover, the strain relief when going from a “sp<sup>2</sup>” to a sp<sup>3</sup> C atom is a driving force for addition and redox reactions.

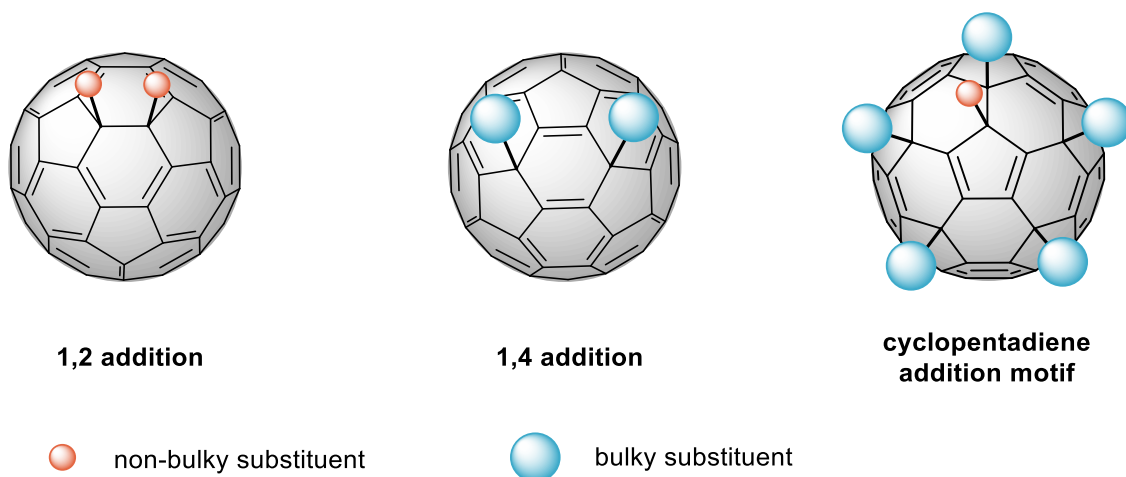
It is also interesting to consider the electronic structures of C<sub>60</sub> in order to explain some structural aspects and the regiochemistry of reactions on C<sub>60</sub>. As seen before in 1.3.1, there is an alternating of bond length. The reason of this alternating is found in the filling of the angular momentum states for an icosahedral structure by the 60 π-electrons. The angular momentum l = 5 level is the last filled level but it is only partially filled by the 10 remaining π-electrons (50 π-electrons completely fill up through l = 4 level and l = 5 level can have up to 22 electrons)<sup>[40,41]</sup>. The frontier orbitals are deduced from the irreducible representation of this l = 5 level, from the lowest to the highest in energy, H<sub>u</sub> + T<sub>1u</sub> + T<sub>2u</sub> for an icosahedral symmetry. The H<sub>u</sub> level which correspond to the HOMO is five-fold degenerated and accept the 10 remaining π-electrons. The T<sub>1u</sub> level is three-fold degenerated and correspond to the LUMO level. It is the filling of HOMO and LUMO levels who have a direct impact on the structure of C<sub>60</sub>. For the HOMO, the bonding interactions are located preferably at the [6,6] junctions and the anti-bonding interactions at the [5,6] junctions<sup>[40]</sup> explaining the bond alternation. As for the low lying LUMO level with anti-bonding interactions at the [6,6] junctions<sup>[40]</sup>, it explained the regiochemistry of addition reactions on C<sub>60</sub> (see 2.2).

### 2.2 Addition and cycloaddition reactions.

The reactivity of C<sub>60</sub> can be assimilated to a strained electron deficient polyolefin and is centered on the [6,6] bonds which can react easily with nucleophilic reagents. C<sub>60</sub> as a good dienophile and dipolarophile can undergo cycloaddition reactions.

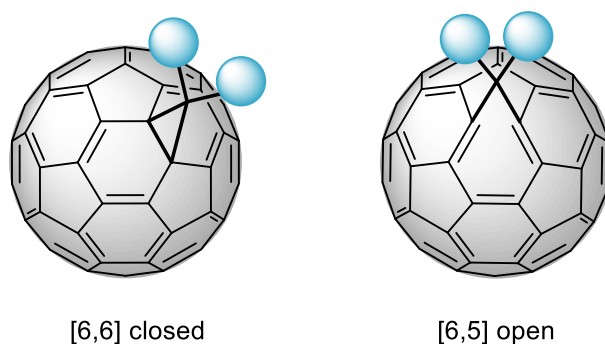
The properties of C<sub>60</sub> have generated a huge interest. However, its poor solubility has limited its utilization. The increase of solubility by functionalizing the C<sub>60</sub> sphere have greatly contributed to the development of fullerene chemistry. In addition, the main properties of the C<sub>60</sub> are kept in mono-adduct derivatives. In this context, a lot of nucleophilic (e.g. Bingel

reactions<sup>[42]</sup>, Grignard reactions<sup>[43]</sup>, Organolithium reactions<sup>[43]</sup>, radical<sup>[44]</sup> and carbene<sup>[45]</sup> additions have been developed in order to functionalize C<sub>60</sub>. For the nucleophilic addition, the nucleophilic intermediate NuC<sub>60</sub><sup>-</sup> can be quenched by the addition of an electrophile E<sup>+</sup>. Addition preferably occurs at 1,2 positions for non-bulky reagents and at 1,4 positions for bulky reagents (Figure I-6).<sup>[46,47]</sup> An original cyclopentadiene motif can be also formed under specific conditions (Figure I-6).<sup>[48-50]</sup>



*Figure I-6 : Different modes of addition of nucleophilic or radical reagents on C<sub>60</sub>.*

A large number of cycloadditions (e.g. Diels-Alder reactions) and dipolar cycloadditions (e.g. azomethine ylides reactions<sup>[51]</sup>, diazo reactions<sup>[52]</sup>) have been also published. Cycloaddition always occurs on the [6,6] bond. Nonetheless in the case of diazo reactions, a rearrangement occurs leading to methanofullerenes with a [6,6]-closed isomerism or a [6,5]-open isomerism (Figure I-7) depending on the reaction conditions.<sup>[52,53]</sup>



*Figure I-7 : Methanofullerenes with a [6,6]-closed isomerism or a [6,5]-open isomerism.*

According to the literature, the Bingel and Diels-Alder reactions are the most used conditions for the multi-functionalization of C<sub>60</sub>. This is one of the reasons why the present manuscript is mainly focused on the Bingel reactions and in a lesser extend to Diels-Alder reactions.

Bingel reactions:

Discovered in 1993 by Carsten Bingel,<sup>[42]</sup> it is a nucleophilic addition of a  $\alpha$ -halomalonate on  $C_{60}$  followed by a nucleophilic intra-molecular substitution that leads to a methanofullerene (Figure I-8). In the initial conditions, a  $\alpha$ -halomalonate was used with sodium hydride (NaH) as base.<sup>[42]</sup> Later some conditions have been developed in order to generate the halogenomalonate *in-situ* with  $I_2$ <sup>[54]</sup> or  $CBr_4$ <sup>[55]</sup> and 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU) as a base. The use of  $I_2$  is more suitable for the synthesis of lower adducts (up to tris-adducts of  $C_{60}$ ) and  $CBr_4$  for higher adducts (e.g. hexa-adducts of  $C_{60}$ ).

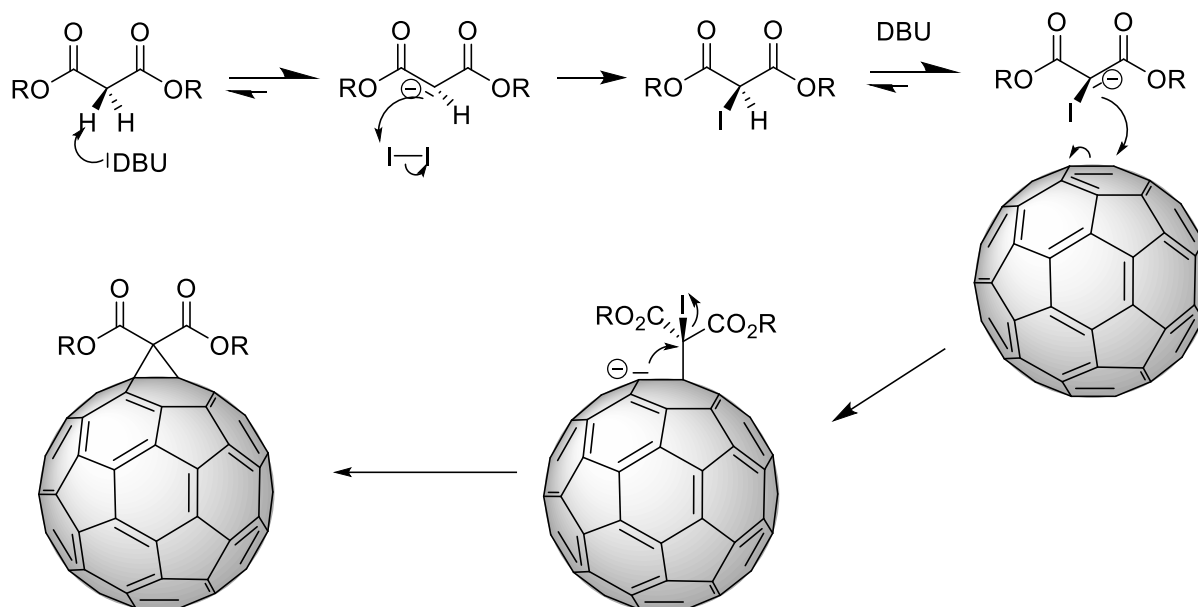


Figure I-8 : Bingel's reaction mechanism with *in-situ* halogenation.

Diels-Alder reactions:

The Diels-Alder reaction is a [4+2] cycloaddition between a diene and a dienophile. The earliest reported characterized products of a Diels-Alder reaction between a diene and  $C_{60}$  was in 1993.<sup>[43,56-58]</sup> An advantage of the use of Diels-Alder reactions for the functionalization of  $C_{60}$  is its reversibility. Diels-Alder adducts can be easily removed by retro-Diels-Alder reactions and thus act as an efficient directing group.<sup>[59]</sup>

## 2.3 Multi-functionalization of $C_{60}$ .

### 2.3.1 Nomenclature.

The 30 double bonds of  $C_{60}$  are all equivalents. Therefore a mono-addition will lead to only one product. The remaining 29 double bonds can be energetically differentiated in to nine different sites. Addition of a second addend to a  $C_{2v}$ -symmetrical  $C_{60}$  mono-adduct can

in principle lead to nine different regioisomeric bis-adducts. Relative to the first addend, the second one can be located either in the same hemisphere (*cis*), in the opposite hemisphere (*trans*) or on the equatorial belt (*e*) (Figure I-9).<sup>[60]</sup> For identical addends, a second attack onto the *e*-edge or *e*-face positions leads to identical products.

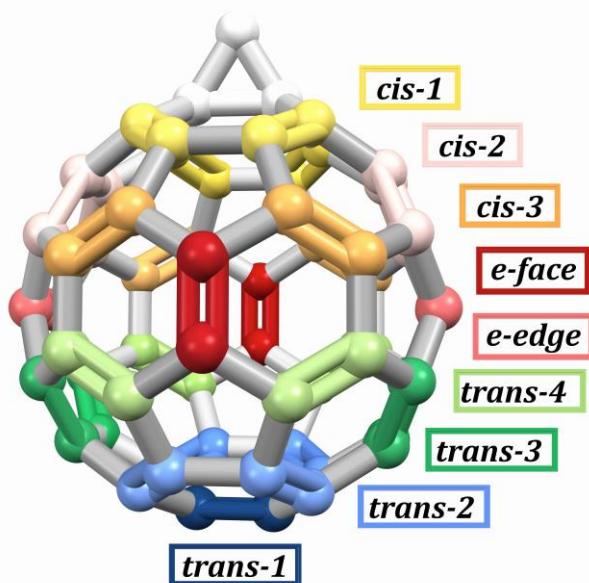


Figure I-9 : Nomenclature of double bonds relative to the first adduct.

In/out descriptor can also be needed, for macrocyclic or dissymmetric compounds, to indicate the direction of the substituents. When considering the substituents A in Figure I-10, there are three possible cases. In case 1 and 3, the two substituents A are together (in/in isomerism) or apart (out/out isomerism). In case 2, the two substituents are mixed (in/out isomerism) considering case 1 and 3.

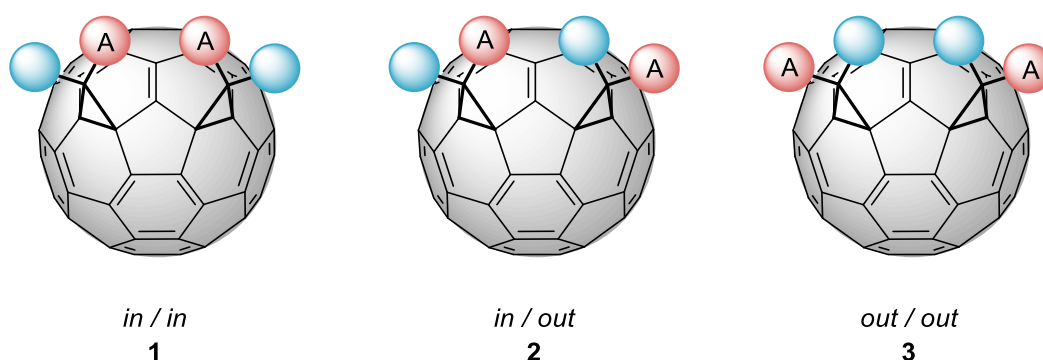


Figure I-10 : Representation of the In/Out Isomerism.

Multiple additions on C<sub>60</sub> can also lead to chiral addition patterns. To indicate the absolute configuration of inherently chiral fullerene, stereodescriptors (<sup>f,s</sup>C) and (<sup>f,s</sup>A) (“f” = fullerene, “s” = systematic numbering, “C” = clockwise, “A” = anti-clockwise) are used. To

determine if it is ( $f^sC$ ) or ( $f^sA$ ), the systematic numbering of the adducts must be the smallest possible (Figure I-11).

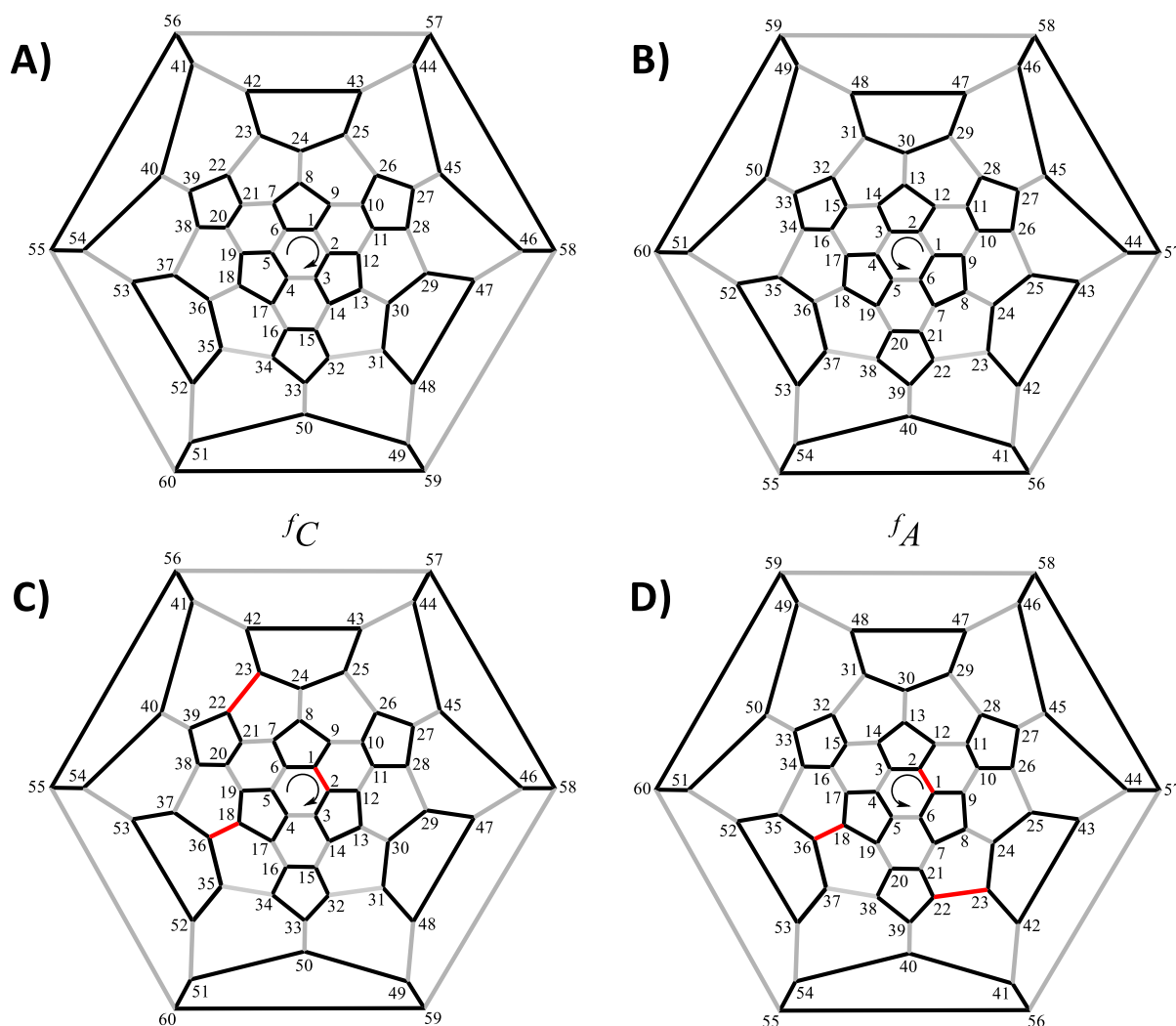


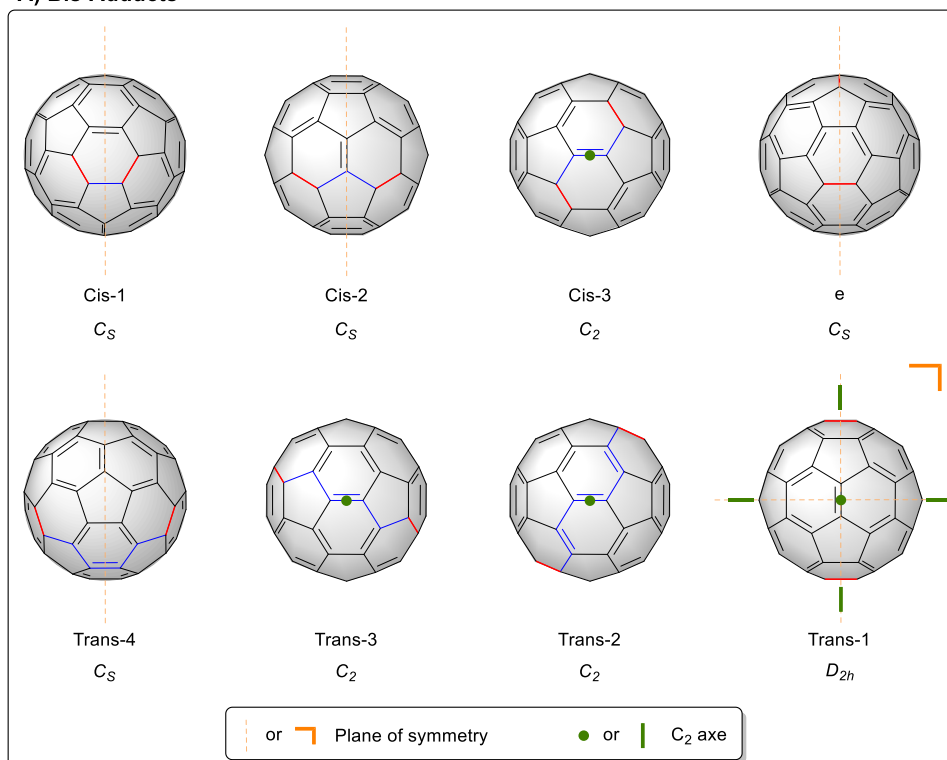
Figure I-11 : Clockwise (A) and Anti-clockwise (B) systematic numbering. C) Representation of ( $f^{-1,2,18,22,23,36}C$ ) tris-adducts. D) Representation of ( $f^{-1,2,18,22,23,36}A$ ) tris-adducts.

### 2.3.2 Symmetry of simple multi-adducts of C<sub>60</sub>.

When multi-adducts of C<sub>60</sub> are synthesized, different regioisomers can be formed. In this context, the symmetry of the formed multi-adduct is an important matter in order to attribute the addition pattern and to characterize each regioisomers. The symmetry of all the possible bis-adducts and selected tris- and hexa-adducts with equivalent malonates are given in Figure I-12.

<sup>13</sup>C NMR analysis can be used to deduce the symmetry of multi-adducts of C<sub>60</sub>. The important signals on <sup>13</sup>C NMR spectrum are signals related to the "C sp<sup>2</sup>" of C<sub>60</sub>.

## A) Bis-Adducts



## B) Tris-Adducts

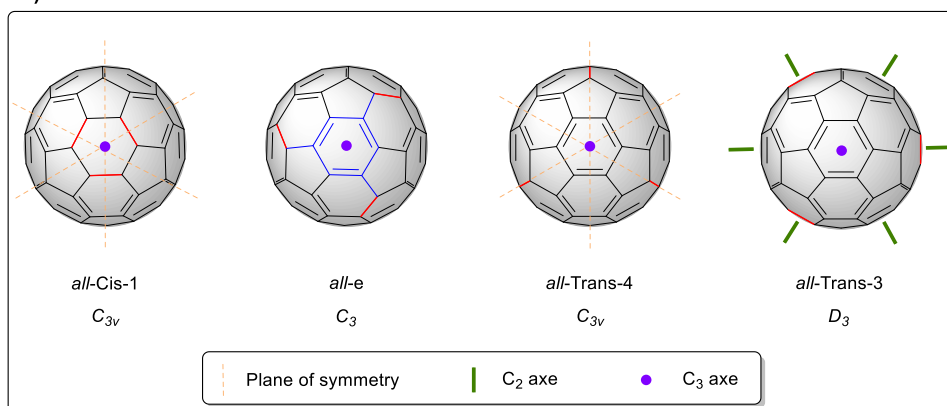
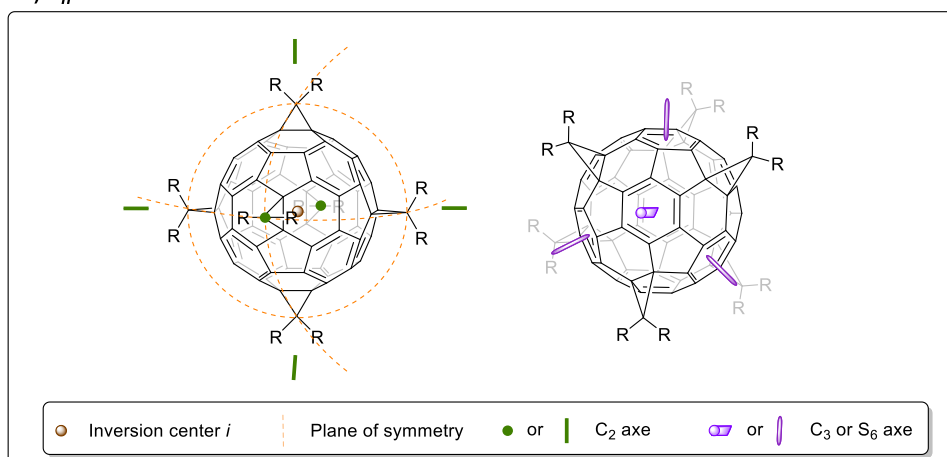
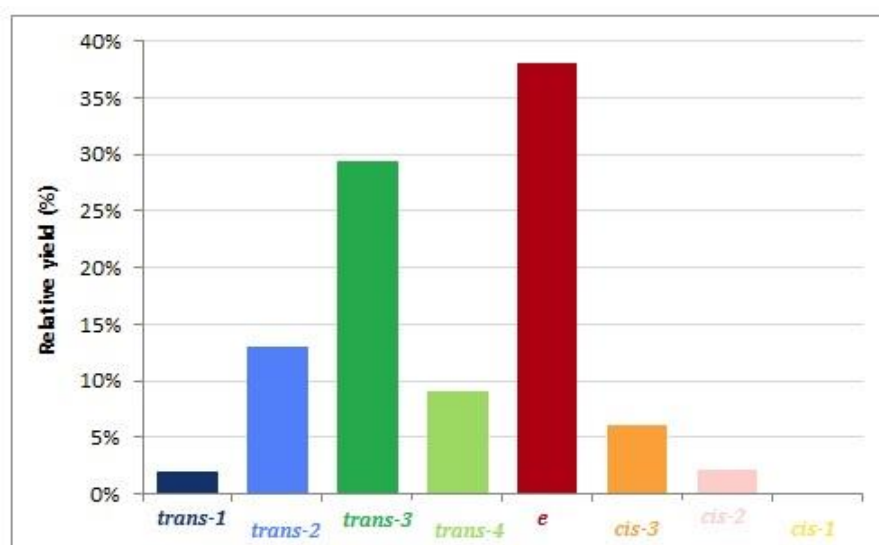
C)  $T_h$ -Hexa-Adducts

Figure I-12 : Representation of the symmetry of all the possible bis-adducts of  $C_{60}$ . Selected examples of tris- and hexa-adducts are also shown.

### 2.3.3 Regiochemistry of twofold additions.

The regiochemistry of twofold additions of malonates on  $C_{60}$  has been studied by Hirsch and coworkers.<sup>[60]</sup> Starting from the mono-adduct  $C_{61}(\text{COOEt})_2$ , they isolated seven out of the eight theoretically possible regioisomers  $C_{62}(\text{COOEt})_4$ . The assignment of each regioisomer was based on both the symmetry deduced from the NMR spectra and the order of elution (polarity) of each product. With the obtained results (Graph 1), an order of reactivity of the double bonds could be established. The *e* double bonds ( $e_{\text{face}}$  and  $e_{\text{edge}}$  are equivalent for the addition of identical symmetrical malonates) are the more reactive sites followed by the *trans*-3 double bonds. In the case of addition of non-identical or dissymmetrical addends, a difference can be made between  $e_{\text{face}}$  and  $e_{\text{edge}}$  double bonds. The  $e_{\text{edge}}$  double bond is slightly more reactive than the  $e_{\text{face}}$  double bonds.<sup>[61,62]</sup> The order of reactivity of the other positions is *trans*-2 > *trans*-4 > *trans*-1 > *cis*-3 > *cis*-2. The *trans*-1 position is more reactive than the *cis*-3 and *cis*-2 positions because one should consider the fact that there is only one *trans*-1 double bond for four double bonds for the *cis*-3 or *cis*-2 positions. Furthermore, *cis* positions are susceptible to steric hindrance effects. This is not the case for *e* and *trans* positions. *Cis*-1 are not really the least reactive sites, in fact, it should be one of the more reactive positions.<sup>[61]</sup> In this particular case *cis*-1 bis-adducts cannot be formed due to steric effects.



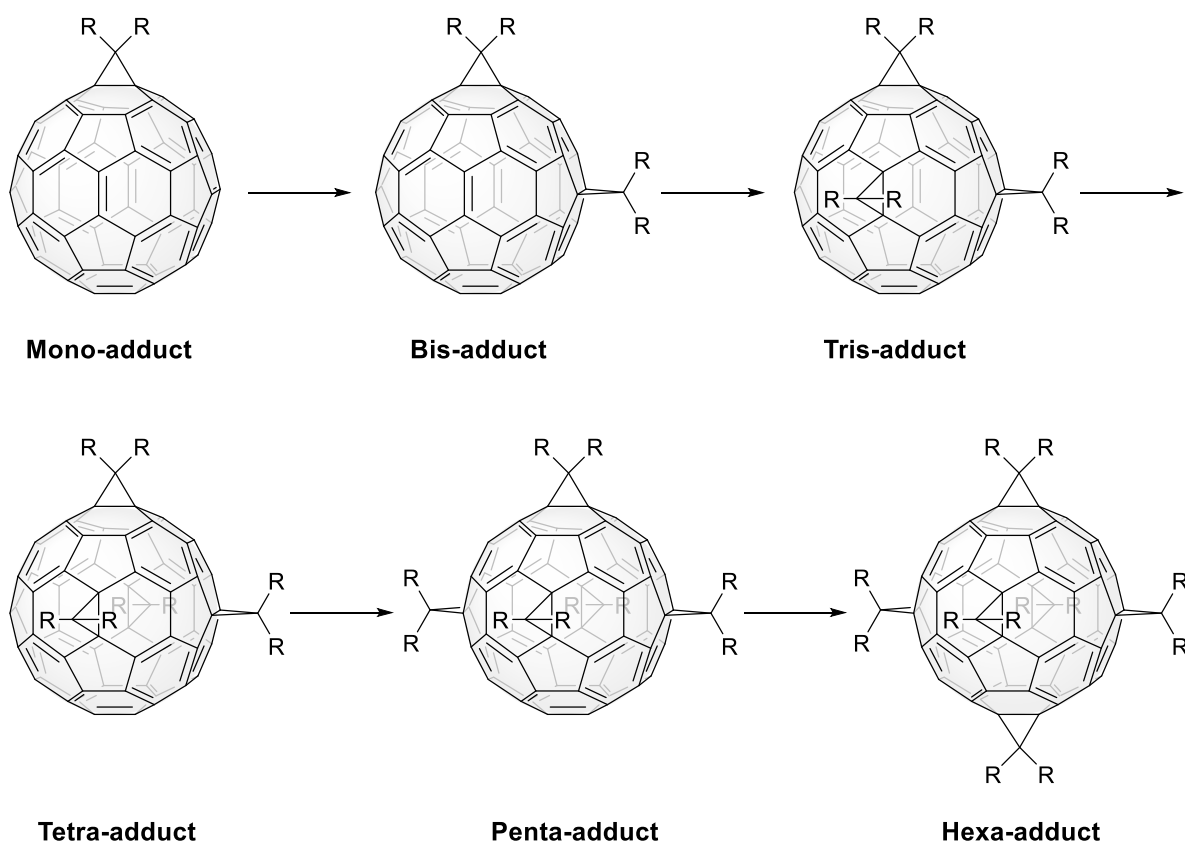
Graph 1 : Relative yields of the different regioisomers.

### 2.3.4 Multiple additions on $C_{60}$ .

The extreme complexity of the multi-fonctionalization of  $C_{60}$ , requires a good understanding of the reactivity of the multi-adducts of  $C_{60}$  and more particularly their regiochemistry. As seen in the previous paragraphs, 9 regioisomers can be formed for a twofold addition. This number increases to 46 regioisomers for a threefold addition. By a step-by-step procedure, the *e,e,e* tris-adducts (each addends are in a *e* position relative to each other) and the *trans*-3, *trans*-3, *trans*-3 tris-adducts, starting from the *e* bis-adducts and the *trans*-3 bis-adducts respectively, were the only ones isolated among the multitude of formed

regioisomers.<sup>[60]</sup> In the continuity of this work, the Hirsch' group has synthesized the  $T_h$ -symmetrical hexa-adducts following the same procedure.<sup>[62]</sup> Starting from the  $e,e,e$  tri-adducts, only two tetra-adducts were formed. For the fifth and sixth additions, only one penta-adducts and hexa-adducts were formed. Based on this step-by-step synthesis of the  $T_h$ -symmetrical hexa-adducts from the mono-adducts (Figure I-13), some important points can be deduced:

- $e$  and  $trans$ -3 positions are the preferred reactions sites.
- The regioselectivity is increasing with the degree of addition. The second and third additions have a weak regioselectivity. The fourth and fifth additions are more regioselective and the sixth is totally regioselective.
- This last point can lead to a general rule that says: "the more addends bound in  $e$  positions to a certain double bond, the more favorable is an attack of this bond".<sup>[62]</sup>
- In opposition to the regioselectivity, the reactivity of the  $C_{60}$  sphere is decreasing with each subsequent addition.



*Figure I-13 : Step-by-step synthesis of the  $T_h$ -symmetrical hexa-adducts from the mono-adducts.*

For the formation of  $T_h$ -symmetrical hexa-adducts in a one-step synthesis, the high reactivity of  $e$  double bonds is definitely an asset but yields are substantially increased with the activation of the  $e$  positions. The first published example of  $T_h$ -hexa-substituted  $C_{60}$  has highlighted the importance of the addition reversibility of the addends for the activation of the  $e$  positions.<sup>[63]</sup> Malone and coworkers were able to obtain a  $T_h$ -hexa-substituted platinum



derivatives  $\{[(C_2H_5)_3P]_2Pt\}_6C_{60}$  with an excellent yield of 88% by reversible multiple additions of a Platinum complex  $[(C_2H_5)_3P]_4Pt$  on  $C_{60}$ . Unfortunately, the addition of malonates on  $C_{60}$  are irreversible. In order to have a relatively good yield for the formation of  $T_h$ -symmetrical hexa-adducts, the activation of e positions must be done by an auxiliary reagent which add reversibly on  $C_{60}$ . In search of such an auxiliary reagent, Hirsch and coworkers have developed conditions with 9,10-dimethylantracene (DMA).<sup>[59]</sup> DMA adds reversibly at room temperature to form Diels-Alder adducts  $C_{60}(DMA)_n$ , acting as templates for the activation of the e positions. Yields are considerably increased with template activation than without.<sup>[55,59]</sup> Even more efficient conditions have been further developed by Sun and coworkers in 2005.<sup>[64]</sup> They modified the conditions developed by Hirsch's group<sup>[55]</sup> with the use of a large excess of  $CBr_4$  (5-10 times more). Under these conditions, they were able to form hexa-adducts from larger malonate reagents. Moreover DMA is not necessary anymore when a large excess of  $CBr_4$  is used,<sup>[65]</sup> because it was shown that  $CBr_3^-$  plays the directing effect owing to its reversible addition onto the fullerene.<sup>[66]</sup>

Formation of  $T_h$ -hexa-adducts are sensitive to steric hindrances. Indeed, when starting from malonates bearing bulky substituents, yields are low or the hexa-adducts are not obtained at all. To resolve this problem, a "click approach" had been proposed (see 3.3).<sup>[67,68]</sup>

The formation of hepta-adducts or higher adducts are not possible with malonates from the  $T_h$ -symmetrical hexa-adducts because all the remaining double bond have a *cis-1* relationship with one already bond malonate. However, it is possible with a suitable addition pattern. Diederich and coworkers have synthesized the  $C_{60}$  hexa-adduct **A** with an original addition pattern (Figure I-14). The further functionalization of **A** led to the octa-adduct **B**.<sup>[69]</sup>

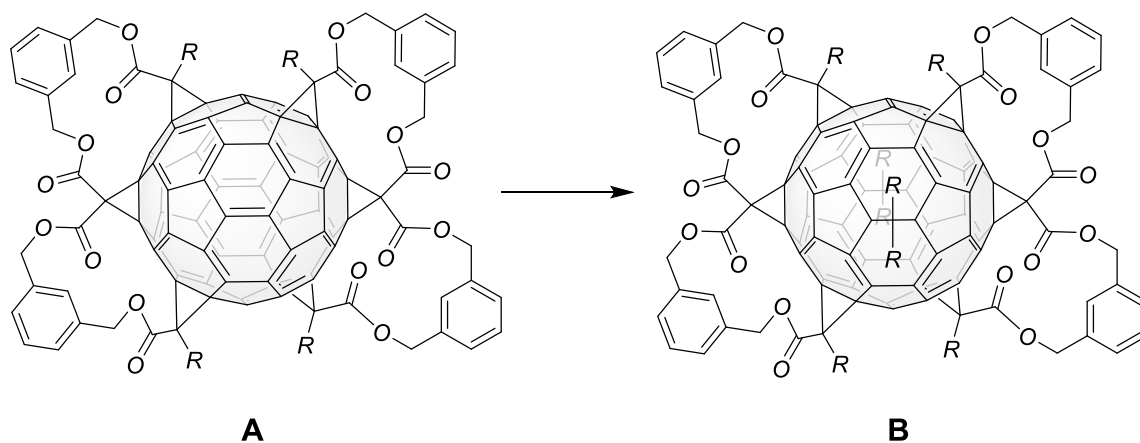


Figure I-14 :  $C_{60}$  hexa-adduct with an original addition pattern and its subsequent octa-adduct.

### 3. The “Click chemistry” in the fullerene chemistry.

#### 3.1 The “click” concept.

The “click” concept was introduced in 2001 by *H. C. Kolb, M. G. Finn* and *K. B. Sharpless*.<sup>[70]</sup> They defined the “click chemistry” with the following set of requirements that reactions must meet:

- *Modular and wide in scope.*
- *Give very high yields.*
- *Generate only inoffensive byproducts.*
- *Be stereospecific.*
- *Have simple reaction conditions.*
- *Readily available starting materials and reagents.*
- *The use of no solvent or a benign solvent or easily removed.*
- *Simple product isolation.*

To meet these requirements, reactions must be energetically favorable with a high thermodynamic driving force. In this context, “click” reactions can be presented as easy, fast and efficient reactions.

A classification<sup>[70,71]</sup> of four families of “click” reactions is possible (Figure I-15):

- Cycloaddition reactions, e.g. 1,3-dipolar cycloaddition and Diels-Alder reactions.
- Nucleophilic ring-opening reactions, such as for epoxides, aziridines, cyclic sulfates, cyclic sulfamidates.
- Carbonyl chemistry of the “non-aldol” type, e.g. the formation of oxime ethers, hydrazones and aromatic heterocycles.
- Additions to carbon-carbon multiple bonds, such as epoxidation, dihydroxylation, aziridination, nitrosyl and sulfenyl halide additions, and some Michael addition reactions.

Since its introduction in 2001, the “click chemistry” has generated a large number of articles, crossing different branches of sciences<sup>[72]</sup> such as materials science or biology. The most famous and most used “click” reaction is certainly the copper catalyzed *Huisgen type* 1,3-dipolar cycloaddition. This reaction has been revealed to be a powerful tool in different fields of chemistry.<sup>[73]</sup>

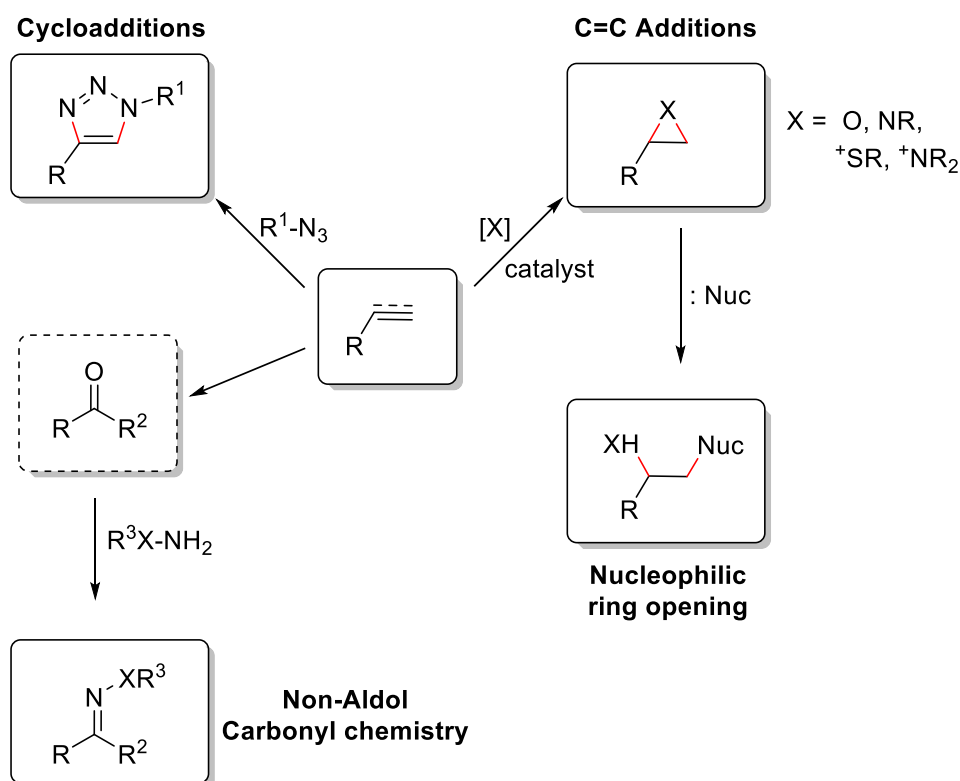


Figure I-15 : Examples of « click » reactions.

### 3.2 The Huisgen type 1,3-dipolar cycloaddition.

The *Huisgen type* 1,3-dipolar cycloaddition corresponds to a cycloaddition reaction between an azide and a terminal alkyne to form a [1,2,3]-triazole. Initially developed by *Rolf Huisgen* in the 60's, a mixture of azides and terminal alkynes was heated at high temperature leading to slow and non-regioselective reactions.<sup>[74,75]</sup> In 2002, *Meldal and coworkers* revisited this reaction with the use of copper(I) salts as catalyst that leads to a fast and regioselective formation at room temperature of the desired [1,2,3]-triazole.<sup>[76]</sup> Later *Sharpless and coworkers* had increased the scope of this new found conditions<sup>[76]</sup> by the use of a more polar solvent such as  $H_2O$  and with the *in situ* reduction of  $Cu^{2+}$  to  $Cu^+$ .<sup>[77]</sup> With the use of copper(I) salts as catalysts, only regioisomers in 1,4 positions are obtained.

#### Mechanism of copper(I) catalyzed 1,3-dipolar cycloaddition.

The first step is a coordination of a terminal alkyne to the copper(I) complex (Figure I-16). That  $\pi$ -complexation lowers the  $pK_a$  of the alkyne C–H by up to 9.8 pH units,<sup>[78]</sup> making possible the formation of copper acetylide in an aqueous media without the addition of a base. In 2013, *Folkin and coworkers* demonstrated that dinuclear copper intermediates are the active species in azide-alkyne cycloadditions.<sup>[79]</sup> The direct observation and isolation of bis(copper) key intermediates was published in 2015 by *Bertrand and coworkers*.<sup>[80]</sup> In the next step, the substituted N-atom coordinates the second copper atom, followed by

nucleophilic attack at the terminal N-atom of the azide by the  $\beta$ -carbon of the acetylide to form the first covalent C–N bond (Figure I-16).<sup>[79]</sup> The ring closure happens next to form a triazolide which is subsequently protonated to form the final triazole substituted in 1,4 positions.

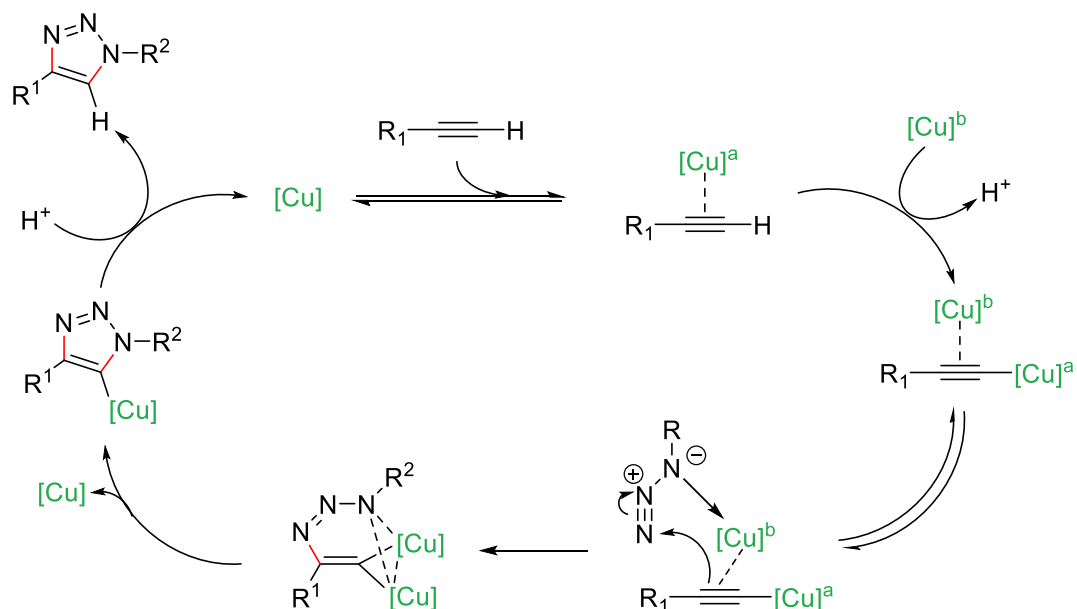


Figure I-16 : Proposed mechanism of copper(I) catalyzed 1,3-dipolar cycloaddition.

### 3.3 The Copper-catalyzed alkyne-azide 1,3-dipolar cycloaddition in Fullerene chemistry.

#### 3.3.1 Introduction.

The copper-catalyzed alkyne-azide 1,3-dipolar cycloaddition (CuAAC) reaction is a powerful tool for the synthesis of a large variety of molecules from small to macromolecular size with good yields.<sup>[73]</sup> On the other hand, direct functionalization of  $C_{60}$  in one and final step can be limited by steric effects, especially for highly substituted  $C_{60}$ .<sup>[81]</sup> Post-functionalization of quite simple  $C_{60}$  building blocks is a solution to form super-functionalized  $C_{60}$  derivatives without steric hindrance problems. With its high tolerance to a large variety of functions and its low sensibility to steric effects, the CuAAC reaction arises as a perfect candidate to considerably increase the scope of post-functionalization of  $C_{60}$  derivatives in respect to other more restricted reactions such as esterifications,<sup>[82,83]</sup> amidifications<sup>[84]</sup> and condensations.<sup>[85,86]</sup>

The first published article of CuAAC reactions in fullerene chemistry revealed the potential of this kind of reaction for the post-functionalization of  $C_{60}$  derivatives.<sup>[67]</sup> Nakamura and coworkers reported the straightforward synthesis of fullerene-carbohydrate conjugates in very good yields by CuAAC reactions between a pentaalkynylfullerene building block and functionalized azides (Figure I-17). Furthermore, they highlighted the mild conditions of

CuAAC reactions that did not require the protection of the sugar hydroxyl groups, facilitating the synthesis.

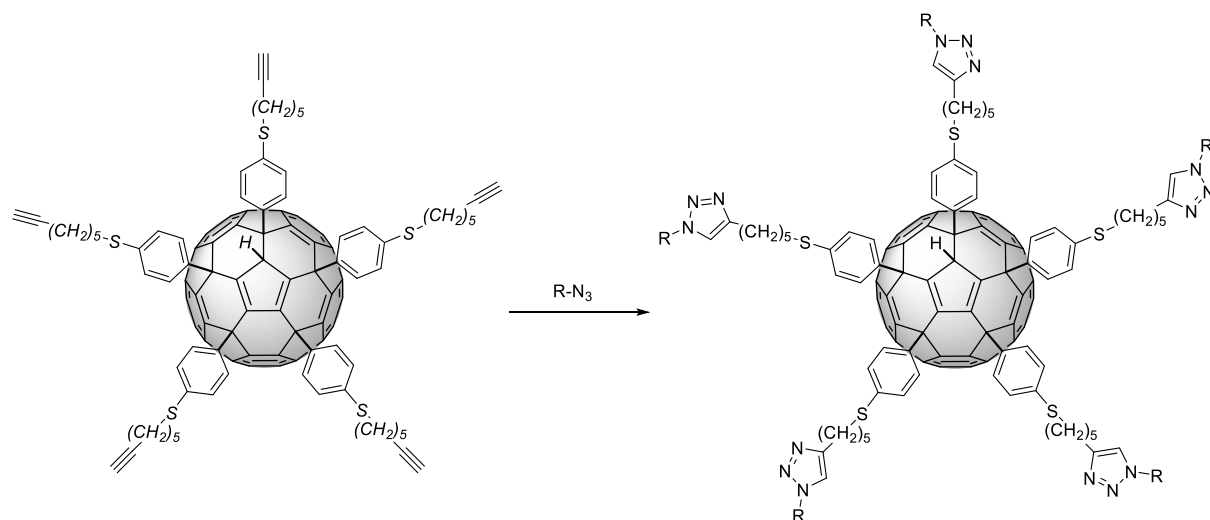


Figure I-17 : First examples of post-functionalization of C<sub>60</sub> derivatives by CuAAC reactions.

The potential of CuAAC reactions on different C<sub>60</sub> derivatives bearing malonates have been investigated in the Nierengarten's group.<sup>[87]</sup> In the following paragraphs, important results and some examples of this particular field are summarized.

### 3.3.2 CuAAC reactions with Mono-, Bis- and Tris-adducts of C<sub>60</sub>.

The elaboration of mono-adducts building blocks (Figure I-18) revealed some limitations for the use of CuAAC reactions in fullerene chemistry. Indeed, organic azides undergo [3+2] cycloadditions with the [6,6] double bonds of fullerenes.<sup>[88,89]</sup> When the mono-adducts bears terminal alkyne groups, the solubility is an important parameter. Indeed, reactions with moderately soluble alkyne-substituted mono-adducts are slow and side reactions are observed decreasing the yield (48% with **I-1**).<sup>[90]</sup> However, when the alkynylated mono-adducts are sufficiently soluble better yields are obtained (80% with **I-2**).<sup>[90,91]</sup> The use of mono-adducts building blocks bearings azide groups is more delicate. These kind of compounds are very unstable in the solid state.<sup>[90]</sup> Fullerene azide (**I-3**) requires to be always in solution to avoid polymerization by intermolecular reaction of azide with C<sub>60</sub>. Under this storage conditions, subsequent CuAAC reactions can be performed with good yields (e.g. 97% with **I-3**).<sup>[92]</sup>



3.3.3 CuAAC reactions with  $T_h$ -addition pattern hexa-adducts.

## 3.3.3.1 Building blocks

The preparation of hexa-adducts with  $T_h$ -addition pattern are sensitive to steric hindrances. Thus, it can be difficult to form highly substituted hexa-adducts by direct additions of bulky malonates. Our group has shown that the post-functionalization of easily accessible hexa-adducts bearing 12 terminal alkyne or azide groups can be easily performed under CuAAC conditions. The hexa-adducts building blocks bearing 12 terminal alkyne (**I-8**) or azide (**I-9**) groups were synthesized with good yields (49% and 62%) starting from malonates **I-6** and **I-7** respectively (Figure I-20).<sup>[68,97]</sup> For **I-8**, the deprotection of trimethylsilyl (TMS) groups can be performed before the CuAAC reaction. Alternatively, the deprotection can be carried out *in situ* during the CuAAC reaction by adding TBAF to the click mixture. Compound **I-9** is enough stable to be stored 2-3 days at low temperature without intra or intermolecular cycloaddition reaction between the azide and the  $C_{60}$  core.

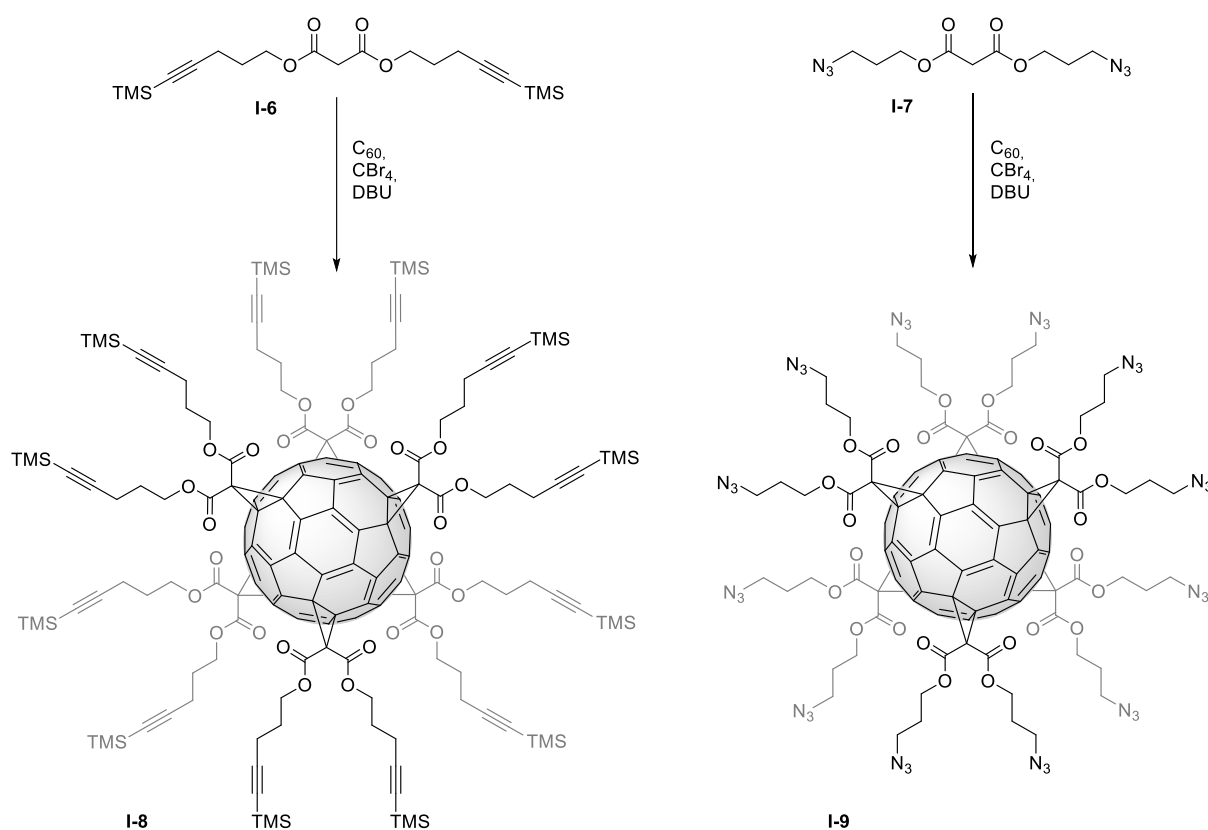
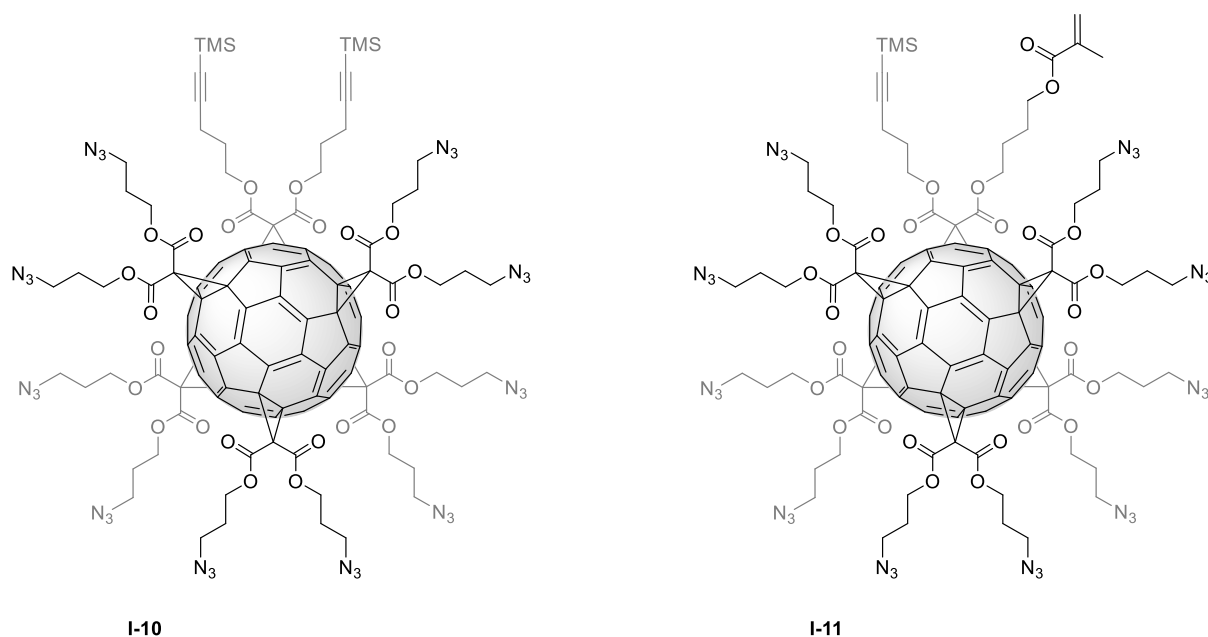


Figure I-20 : Classical  $T_h$ -symmetrical hexa-adducts building blocks.

Another example of  $T_h$ -symmetrical hexa-adducts has been reported by Bräse and coworkers.<sup>[98]</sup> In this particular case, the use of macrocyclic malonates bearing an azide function gave rise to hexa-adduct building blocks allowing for the grafting of six peripheral subunits.

The two last examples of reported building blocks with  $T_h$ -addition pattern are mixed [5,1] hexa-adducts **I-10** and **I-11** (five malonates are identical and one malonate is

different)(Figure I-21). The particularity of I-11 is that three successive post-functionalization reactions can be made (two by CuAAC reactions and one by radical thiol-ene click reaction).



*Figure I-21 : Mixed [5,1] hexa-adducts building blocks.*

The advantage of this methodology is that the building block can be adequately chosen in order to simplify the synthesis of the “clickable” functions in addition to the formation of highly functionalized hexa-adducts.

### 3.3.3.2 Applications in various fields.

Fullerene hexa-adducts have been used in various fields. In this section, only a brief overview is given for each application.

All the clicked fullerene hexa-adduct derivatives described in this section were obtained in good yields from  $\approx 60\%$  to  $90\%$ . For hexa-adducts, electrons or energy transfers on the  $C_{60}$  core are very weak.<sup>[99,100]</sup> In general, the  $C_{60}$  hexa-adducts core plays only the role of a scaffold.

#### Photoactive clicked fullerene derivatives:

Three examples of photoactive derivatives have been published by Nierengarten’s group and collaborators.

The interest of hexa-substituted fullerene derivative for photoactive applications is related to the perturbation of both ground- and excited-state energy levels of the carbon sphere. Therefore, the absorption spectra of the central core is blue-shifted and the potential of the first reduction wave is shifted to about 1 V.



Starting from the building block **I-9**, hexa-adducts **I-12**<sup>[68]</sup> and **I-13** were prepared (Figure I-22). Compound **I-12** has a very strong absorption at 420 nm (Soret band, molar absorptivity:  $431 \times 10^4 \text{ M}^{-1} \cdot \text{cm}^{-1}$ ) owing to the 12 peripheral Zn<sup>II</sup>-porphyrin. Photoinduced electron transfer and singlet-singlet energy transfer between the porphyrin and the fullerene core are prevented by the unfavorable electronic states. Thus, **I-12** is attractive for the construction of a supramolecular photosynthetic model by association with a C<sub>60</sub>-imidazole mono-adducts derivative.<sup>[100]</sup> In contrast, when hexa-adducts are surrounded by blue emitter chromophores, photoinduced energy transfer to the core can be possible which is not the case for redshifted chromophores. Compound **I-13** bearing 12 stilbene sub-units exhibits efficient light-harvesting properties from the periphery to the core by an efficient singlet-singlet energy transfer.<sup>[99]</sup>

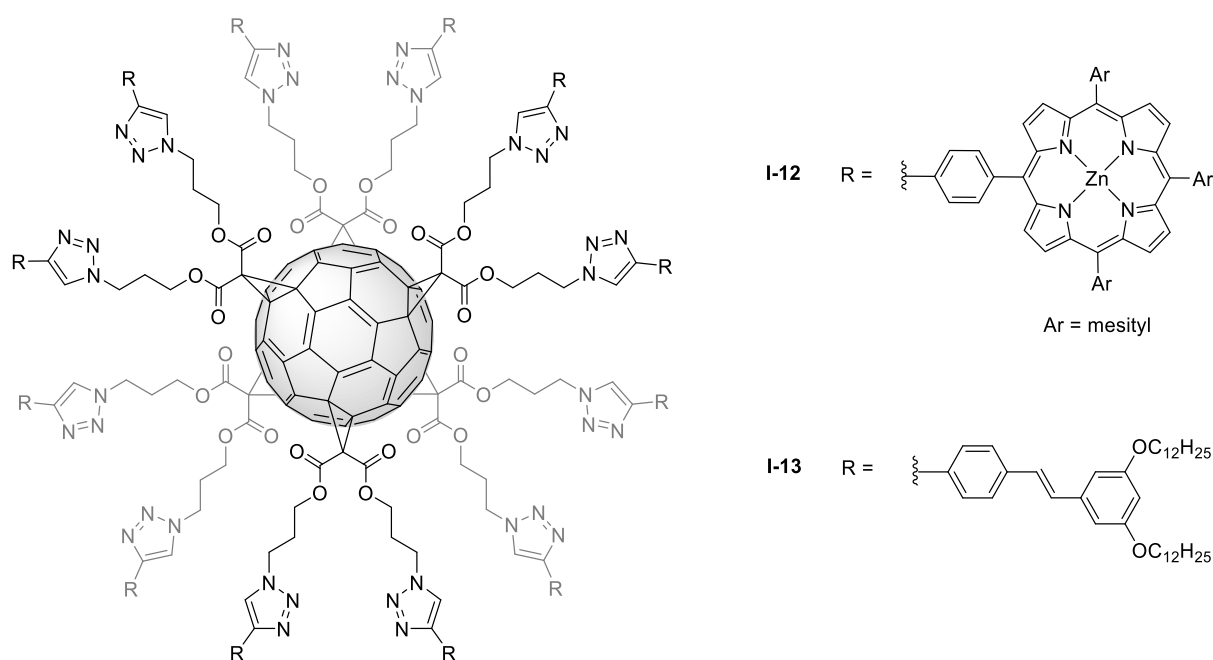


Figure I-22 : Examples of photoactive clicked hexa-adducts.

In the last example,<sup>[99]</sup> building block **I-10** was used in order to associate ten yellow dyes with two complementary blue dyes to form **I-14** (Figure I-23). Photophysical studies of **I-14** has shown energy transfers from the yellow to the blue dyes resulting to a stronger fluorescence of the blue dyes. **I-14** can be considered as a solar energy concentrator.

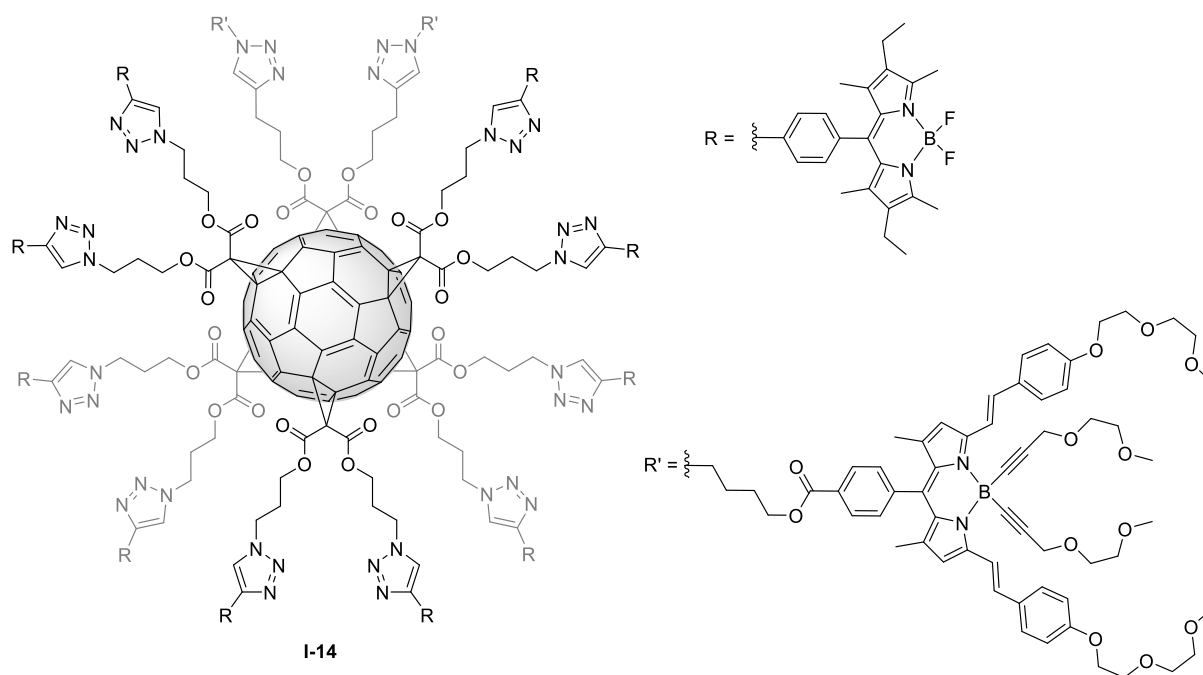


Figure I-23 : Example of photoactive clicked mixed [5,1] hexa-adducts.

#### Electroactive clicked fullerene derivatives:

In contrast to  $C_{60}$ , fullerene hexa-adducts with an octahedral addition pattern are very poor electron acceptors. The  $C_{60}$  core of the hexa-adducts is electrochemically silent over a large potential window (from ca. -1.5 to +1.5 V vs. a standard calomel electrode (SCE)). Therefore, the electrochemical properties of fullerene hexa-adduct derivatives are directly related to the substituents.

Molecular motion of **I-15** (Figure I-24) by electrochemical control has been reported.<sup>[101]</sup> The molecular motion is induced by redox reactions on viologen subunits which  $\pi$ -dimerize when radical cations are produced.

**I-16** was functionalized with ten ferrocene and two 1,2-dithiolane units. The dithiolane units served as anchors onto a gold surface (Figure I-24). The electrochemical behavior of **I-16** was investigated by cyclic voltammetry. Depending on the scan rate, all or a fraction of the ferrocene subunits can be oxidized.<sup>[102]</sup>

#### Liquid crystalline clicked fullerene derivatives:

Different Percec-type dendrons were grafted onto  $T_h$ -symmetrical  $C_{60}$  hexa-adduct building blocks (Figure I-25). Mesomorphic properties were evidenced by Differential Scanning Calorimetry (DSC) and Polarized Optical Microscopy (POM) measurements. These dendronized molecules self-assemble into unprecedented supramolecular discs containing the fullerene at their core. The addition pattern have little if any effects on the liquid crystal phase.<sup>[103]</sup>

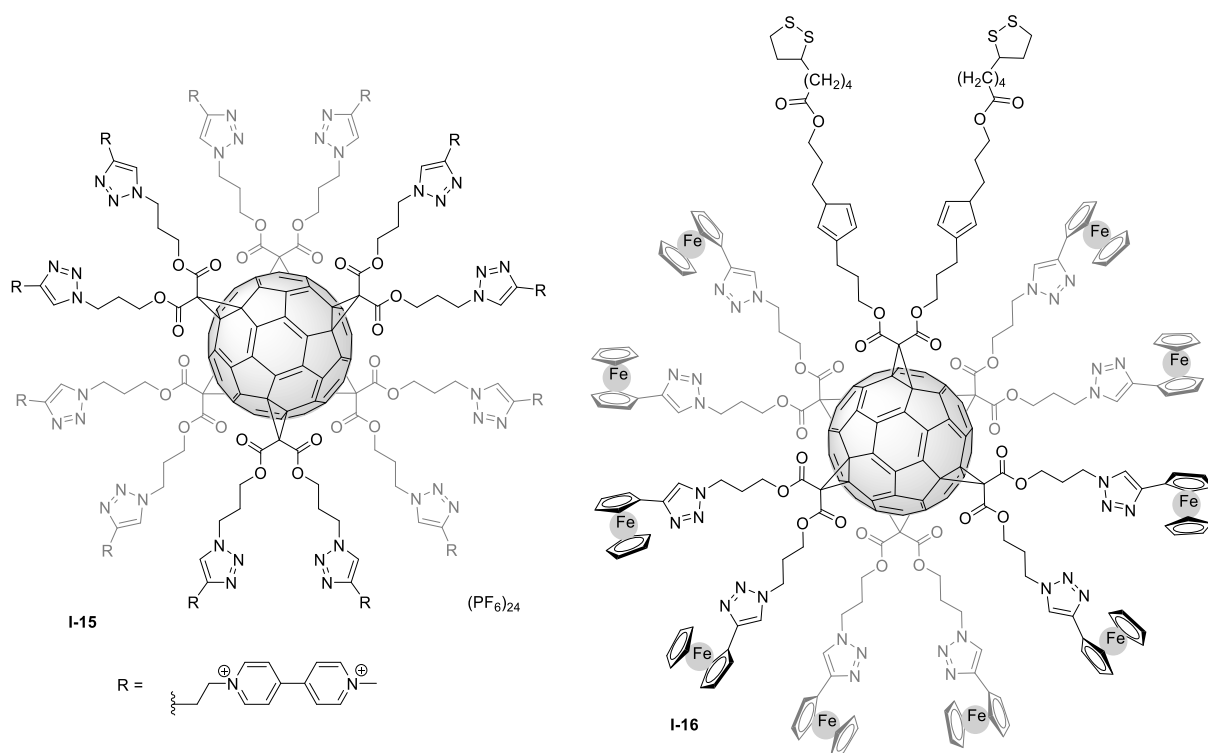


Figure I-24 : Examples of electroactive clicked fullerene derivatives.

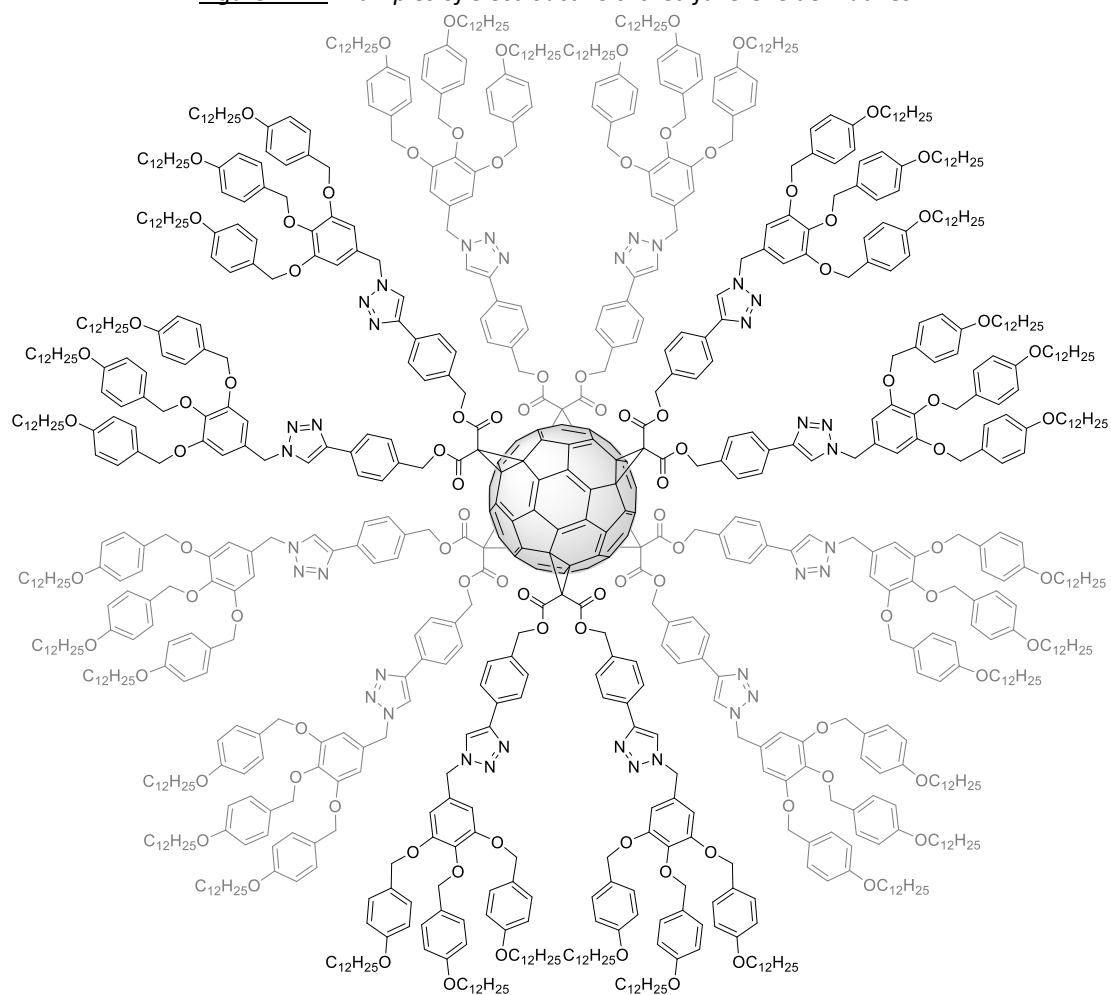


Figure I-25 : Example of a liquid crystal fullerene hexa-adduct derivative.

Bioactive clicked fullerene derivatives:

$T_h$ -symmetrical fullerene hexa-adduct is an interesting scaffold for biological applications. The globular shape is an asset to prevent amphiphilic character and thus possible aggregation in aqueous media. Furthermore, the peripheral function multiplicity of 12 provides a fast dendritic growth.

Preparation of dendronized polycationic hexa-adducts of  $C_{60}$  has shown remarkable gene-delivery capabilities (Figure I-26). These compounds exhibit a low toxicity. The polycationic system is more compact with the globular shape of  $T_h$ -hexa-adducts. Thus, the efficiency can be good even for low generation dendrons.<sup>[104]</sup>

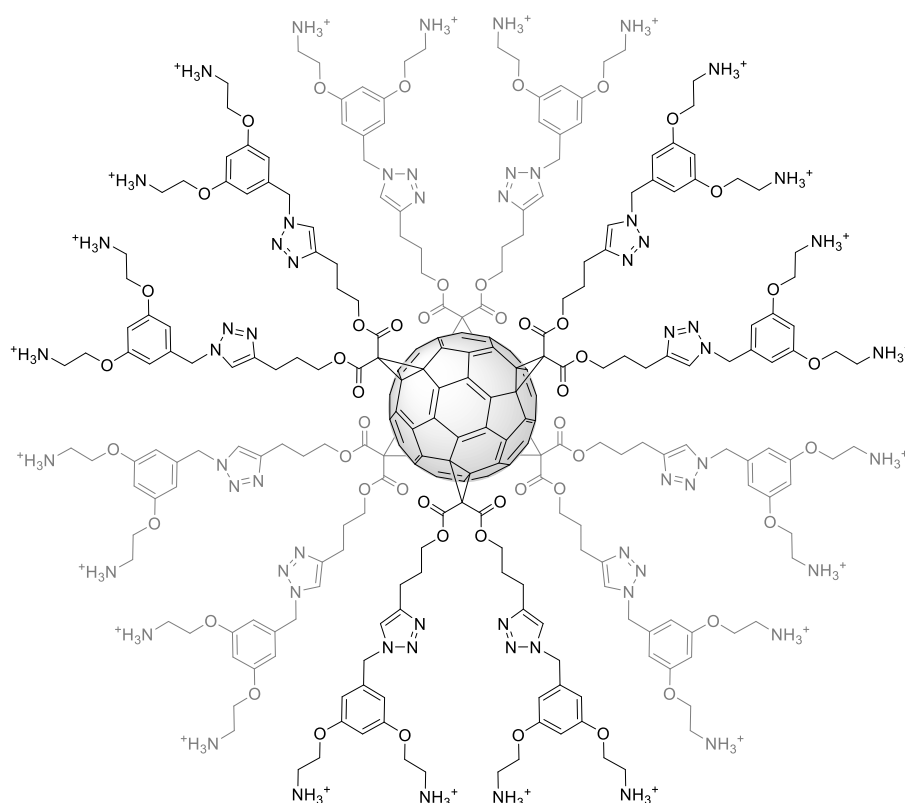


Figure I-26 : Example of a dendronized polycationic hexa-adduct of  $C_{60}$ .

Several hexa-adducts of  $C_{60}$  bearing sugar units at its periphery (Figure I-27) have been reported.<sup>[105–110]</sup> Preparation of glycofullerenes are simplified with the use of CuAAC reactions. These fullerenes derivatives have a good solubility in aqueous media and low cytotoxicity. Multivalent effects were observed for enzymatic inhibition.

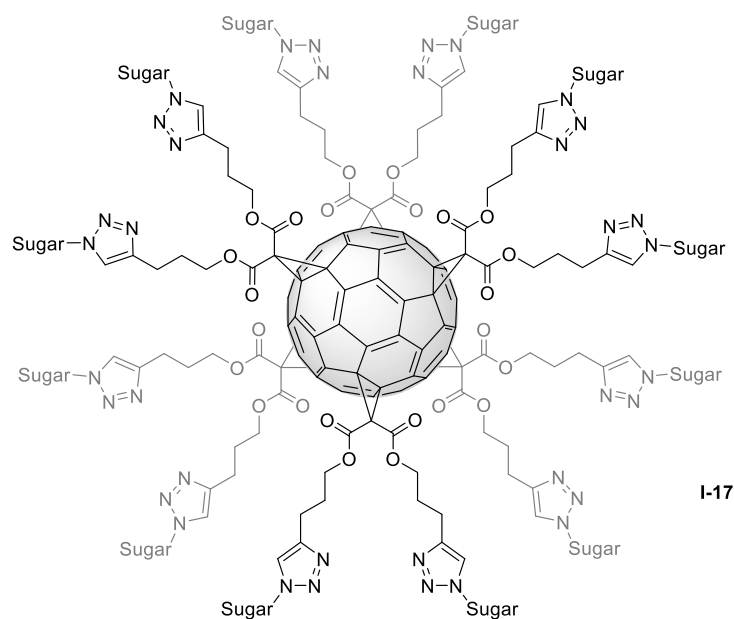


Figure I-27 : Example of bioactive clicked hexa-adducts.

## 4. Objectives.

Fullerene hexa-adducts with a  $T_h$ -symmetrical octahedral addition pattern are a unique class of three dimensional molecules. Their spherical framework is indeed an appealing platform for the preparation of new nanomaterials and bioactive molecules. The direct synthesis of fullerene hexa-adducts from C<sub>60</sub> and malonates is however difficult and generally restricted to relatively simple malonate derivatives thus limiting their accessibility. This major problem was recently solved by producing easily accessible C<sub>60</sub> hexa-adduct derivatives bearing 12 terminal groups allowing their further functionalization to generate structurally more complicated systems. Among them, the copper mediated Huisgen 1,3-dipolar cycloaddition of azides and alkynes resulting in 1,2,3-triazoles was found to be particularly interesting as it allows for the incorporation of almost any functional groups around the fullerene core. The development of an efficient toolbox for the synthesis of fullerene hexa-adducts bearing two or more different peripheral functional subunits is now one of the major challenge to produce a unique multifunctional nanomaterials platform.

The aim of this PhD thesis is to develop in a first step a regioselective methodology for the synthesis of bis- and tris-adducts of C<sub>60</sub>, and in a second step to develop versatile fullerene hexa-adduct building blocks incorporating complementary reactive centers with different and selective reactivity.

The **Chapters II** and **III** were dedicated to the regioselective bis- and tris-functionalization of C<sub>60</sub> respectively by a macrocyclic approach and by a tether-directed approach. Based on the regioselective syntheses of multi-adducts of C<sub>60</sub>, the preparation of mixed C<sub>60</sub> hexa-adducts building blocks and their subsequent post-functionalization were performed in the **Chapter IV**. In the **Chapter V**, the mixed C<sub>60</sub> hexa-adducts building blocks previously prepared are used for the preparation of original fullerodendrimers.

## 5. Bibliography.

- [1] D. E. H. Jones, *New Sci.* **1966**, *31*, 493–496.
- [2] W. E. Barth, R. G. Lawton, *J. Am. Chem. Soc.* **1966**, *88*, 380–381.
- [3] E. Osawa, *Kagaku* **1970**, *25*, 854–863.
- [4] E. A. Rohlfing, D. M. Cox, A. Kaldor, *J. Chem. Phys.* **1984**, *81*, 3322–3330.
- [5] H. W. Kroto, J. R. Heath, S. C. O'Brien, R. F. Curl, R. E. Smalley, *Nature* **1985**, *318*, 162–163.
- [6] W. Krätschmer, L. D. Lamb, K. Fostiropoulos, D. R. Huffman, *Nature* **1990**, *347*, 354–358.
- [7] W. H. Powell, F. Cozzi, G. P. Moss, C. Thilgen, R. J.-R. Hwu, A. Yerin, *Pure Appl. Chem.* **2002**, *74*, 629–695.
- [8] R. E. Haufler, J. Conceicao, L. P. F. Chibante, Y. Chai, N. E. Byrne, S. Flanagan, M. M. Haley, S. C. O'Brien, C. Pan, . et al., *J. Phys. Chem.* **1990**, *94*, 8634–8636.
- [9] L. P. F. Chibante, A. Thess, J. M. Alford, M. D. Diener, R. E. Smalley, *J. Phys. Chem.* **1993**, *97*, 8696–8700.
- [10] G. Peters, M. Jansen, *Angew. Chem. Int. Ed. Engl.* **1992**, *31*, 223–224.
- [11] P. Gerhardt, K. H. Homann, *J. Phys. Chem.* **1990**, *94*, 5381–5391.
- [12] J. B. Howard, J. T. McKinnon, Y. Makarovskiy, A. L. Lafleur, M. E. Johnson, *Nature* **1991**, *352*, 139–141.
- [13] C. J. Pope, J. A. Marr, J. B. Howard, *J. Phys. Chem.* **1993**, *97*, 11001–11013.
- [14] P. Gerhardt, S. Löffler, K. H. Homann, *Chem. Phys. Lett.* **1987**, *137*, 306–310.
- [15] C. Yeretzyan, J. B. Wiley, K. Holczer, T. Su, S. Nguyen, R. B. Kaner, R. L. Whetten, *J. Phys. Chem.* **1993**, *97*, 10097–10101.
- [16] J. Théobald, M. Perrut, J.-V. Weber, E. Millon, J.-F. Muller, *Sep. Sci. Technol.* **1995**, *30*, 2783–2819.
- [17] H. Ajie, M. M. Alvarez, S. J. Anz, R. D. Beck, F. Diederich, K. Fostiropoulos, D. R. Huffman, W. Krätschmer, Y. Rubin, . et al., *J. Phys. Chem.* **1990**, *94*, 8630–8633.
- [18] R. Taylor, J. P. Hare, A. K. Abdul-Sada, H. W. Kroto, *J. Chem. Soc. Chem. Commun.* **1990**, 1423–1425.
- [19] K. Chatterjee, D. H. Parker, P. Wurz, K. R. Lykke, D. M. Gruen, L. M. Stock, *J. Org. Chem.* **1992**, *57*, 3253–3254.
- [20] K. C. Khemani, M. Prato, F. Wudl, *J. Org. Chem.* **1992**, *57*, 3254–3256.
- [21] W. A. Scrivens, P. V. Bedworth, J. M. Tour, *J. Am. Chem. Soc.* **1992**, *114*, 7917–7919.
- [22] R. J. Doome, A. Fonseca, H. Richter, J. B. Nagy, P. A. Thiry, A. A. Lucas, *J. Phys. Chem. Solids* **1997**, *58*, 1839–1843.
- [23] T. G. Schmalz, W. A. Seitz, D. J. Klein, G. E. Hite, *Chem. Phys. Lett.* **1986**, *130*, 203–207.
- [24] H. W. Kroto, *Nature* **1987**, *329*, 529–531.
- [25] H. P. Lüthi, J. Almlöf, *Chem. Phys. Lett.* **1987**, *135*, 357–360.
- [26] J. M. Schulman, R. L. Disch, M. A. Miller, R. C. Peck, *Chem. Phys. Lett.* **1987**, *141*, 45–48.
- [27] R. D. Johnson, G. Meijer, D. S. Bethune, *J. Am. Chem. Soc.* **1990**, *112*, 8983–8984.

- [28] W. I. F. David, R. M. Ibberson, J. C. Matthewman, K. Prassides, T. J. S. Dennis, J. P. Hare, H. W. Kroto, R. Taylor, D. R. M. Walton, *Nature* **1991**, *353*, 147–149.
- [29] S. Liu, Y.-J. Lu, M. M. Kappes, J. A. Ibers, *Science* **1991**, *254*, 408–410.
- [30] K. Hedberg, L. Hedberg, D. S. Bethune, C. A. Brown, H. C. Dorn, R. D. Johnson, M. D. Vries, *Science* **1991**, *254*, 410–412.
- [31] R. S. Ruoff, D. S. Tse, R. Malhotra, D. C. Lorents, *J. Phys. Chem.* **1993**, *97*, 3379–3383.
- [32] J. W. Arbogast, A. P. Darmanyan, C. S. Foote, F. N. Diederich, R. L. Whetten, Y. Rubin, M. M. Alvarez, S. J. Anz, *J. Phys. Chem.* **1991**, *95*, 11–12.
- [33] D. Kim, M. Lee, Y. D. Suh, S. K. Kim, *J. Am. Chem. Soc.* **1992**, *114*, 4429–4430.
- [34] D. M. Guldi, *Chem. Commun.* **2000**, 321–327.
- [35] G. Accorsi, N. Armaroli, *J. Phys. Chem. C* **2010**, *114*, 1385–1403.
- [36] Z. Markovic, V. Trajkovic, *Biomaterials* **2008**, *29*, 3561–3573.
- [37] R. C. Haddon, L. E. Brus, K. Raghavachari, *Chem. Phys. Lett.* **1986**, *125*, 459–464.
- [38] Q. Xie, E. Perez-Cordero, L. Echegoyen, *J. Am. Chem. Soc.* **1992**, *114*, 3978–3980.
- [39] C. Bruno, I. Doubitski, M. Marcaccio, F. Paolucci, D. Paolucci, A. Zaopo, *J. Am. Chem. Soc.* **2003**, *125*, 15738–15739.
- [40] A. Hirsch, in *Fuller. Relat. Struct.* (Ed.: P.D.A. Hirsch), Springer Berlin Heidelberg, **1999**, pp. 1–65.
- [41] M. S. Dresselhaus, G. Dresselhaus, P. C. Eklund, in *Sci. Fuller. Carbon Nanotub.*, Academic Press, San Diego, **1996**, pp. 413–463.
- [42] C. Bingel, *Chem. Ber.* **1993**, *126*, 1957–1959.
- [43] F. Wudl, A. Hirsch, K. C. Khemani, T. Suzuki, P.-M. Allemand, A. Koch, H. Eckert, G. Srdanov, H. M. Webb, in *Fullerenes*, American Chemical Society, **1992**, pp. 161–175.
- [44] M. D. Tzirakis, M. Orfanopoulos, *Chem. Rev.* **2013**, *113*, 5262–5321.
- [45] M. Yamada, T. Akasaka, S. Nagase, *Chem. Rev.* **2013**, *113*, 7209–7264.
- [46] D. A. Dixon, N. Matsuzawa, T. Fukunaga, F. N. Tebbe, *J. Phys. Chem.* **1992**, *96*, 6107–6110.
- [47] N. Matsuzawa, T. Fukunaga, D. A. Dixon, *J. Phys. Chem.* **1992**, *96*, 10747–10756.
- [48] P. J. Krusic, E. Wasserman, P. N. Keizer, J. R. Morton, K. F. Preston, *Science* **1991**, *254*, 1183–1185.
- [49] P. R. Birkett, P. B. Hitchcock, H. W. Kroto, R. Taylor, D. R. M. Walton, *Nature* **1992**, *357*, 479–481.
- [50] M. Sawamura, H. Iikura, E. Nakamura, *J. Am. Chem. Soc.* **1996**, *118*, 12850–12851.
- [51] M. Maggini, G. Scorrano, M. Prato, *J. Am. Chem. Soc.* **1993**, *115*, 9798–9799.
- [52] T. SuzuKI, Q. Li, K. C. Khemani, F. Wudl, Ö. Almarsson, *Science* **1991**, *254*, 1186–1188.
- [53] F. Diederich, L. Isaacs, D. Philp, *J. Chem. Soc. Perkin Trans. 2* **1994**, 391–394.
- [54] J.-F. Nierengarten, T. Habicher, R. Kessinger, F. Cardullo, F. Diederich, V. Gramlich, J.-P. Gisselbrecht, C. Boudon, M. Gross, *Helv. Chim. Acta* **1997**, *80*, 2238–2276.
- [55] X. Camps, A. Hirsch, *J. Chem. Soc. [Perkin 1]* **1997**, 1595–1596.
- [56] Y. Rubin, S. Khan, D. I. Freedberg, C. Yeretziyan, *J. Am. Chem. Soc.* **1993**, *115*, 344–345.
- [57] P. Belik, A. Gügel, J. Spickermann, K. Müllen, *Angew. Chem. Int. Ed. Engl.* **1993**, *32*, 78–80.

- [58] M. Tsuda, T. Ishida, T. Nogami, S. Kurono, M. Ohashi, *J. Chem. Soc. Chem. Commun.* **1993**, 1296–1298.
- [59] I. Lamparth, C. Maichle–Mössmer, A. Hirsch, *Angew. Chem. Int. Ed. Engl.* **1995**, *34*, 1607–1609.
- [60] A. Hirsch, I. Lamparth, H. R. Karfunkel, *Angew. Chem. Int. Ed. Engl.* **1994**, *33*, 437–438.
- [61] F. Djojo, A. Herzog, I. Lamparth, F. Hampel, A. Hirsch, *Chem. – Eur. J.* **1996**, *2*, 1537–1547.
- [62] A. Hirsch, I. Lamparth, T. Groesser, H. R. Karfunkel, *J. Am. Chem. Soc.* **1994**, *116*, 9385–9386.
- [63] P. J. Fagan, J. C. Calabrese, B. Malone, *J. Am. Chem. Soc.* **1991**, *113*, 9408–9409.
- [64] H. Li, A. Kitaygorodskiy, R. A. Carino, Y.-P. Sun, *Org. Lett.* **2005**, *7*, 859–861.
- [65] H. Li, S. A. Haque, A. Kitaygorodskiy, M. J. Meziani, M. Torres-Castillo, Y.-P. Sun, *Org. Lett.* **2006**, *8*, 5641–5643.
- [66] J. Iehl, *La chimie “Click” appliquée À Des Dérivés de C60*, Thèse, Université De Strasbourg, **2010**.
- [67] H. Isobe, K. Cho, N. Solin, D. B. Werz, P. H. Seeberger, E. Nakamura, *Org. Lett.* **2007**, *9*, 4611–4614.
- [68] J. Iehl, R. P. de Freitas, B. Delavaux-Nicot, J.-F. Nierengarten, *Chem. Commun.* **2008**, 2450–2452.
- [69] C. R. Woods, J.-P. Bourgeois, P. Seiler, F. Diederich, *Angew. Chem. Int. Ed.* **2000**, *39*, 3813–3816.
- [70] H. C. Kolb, M. G. Finn, K. B. Sharpless, *Angew. Chem. Int. Ed.* **2001**, *40*, 2004–2021.
- [71] H. C. Kolb, K. B. Sharpless, *Drug Discov. Today* **2003**, *8*, 1128–1137.
- [72] J. E. Moses, A. D. Moorhouse, *Chem. Soc. Rev.* **2007**, *36*, 1249–1262.
- [73] J. Lahann, Ed. , in *Click Chem. Biotechnol. Mater. Sci.*, John Wiley & Sons, Ltd, **2009**, pp. i–xix.
- [74] R. Huisgen, *Angew. Chem. Int. Ed. Engl.* **1963**, *2*, 565–598.
- [75] R. Huisgen, G. Szeimies, L. Möbius, *Chem. Ber.* **1967**, *100*, 2494–2507.
- [76] C. W. Tornøe, C. Christensen, M. Meldal, *J. Org. Chem.* **2002**, *67*, 3057–3064.
- [77] V. V. Rostovtsev, L. G. Green, V. V. Fokin, K. B. Sharpless, *Angew. Chem. Int. Ed.* **2002**, *41*, 2596–2599.
- [78] F. Himo, T. Lovell, R. Hilgraf, V. V. Rostovtsev, L. Noodleman, K. B. Sharpless, V. V. Fokin, *J. Am. Chem. Soc.* **2005**, *127*, 210–216.
- [79] B. T. Worrell, J. A. Malik, V. V. Fokin, *Science* **2013**, *340*, 457–460.
- [80] L. Jin, D. R. Tolentino, M. Melaimi, G. Bertrand, *Sci. Adv.* **2015**, *1*, e1500304.
- [81] A. Hirsch, O. Vostrowsky, *Eur. J. Org. Chem.* **2001**, *2001*, 829–848.
- [82] A. Gégout, M. Holler, T. M. Figueira-Duarte, J.-F. Nierengarten, *Eur. J. Org. Chem.* **2008**, *2008*, 3627–3634.
- [83] U. Hahn, J.-F. Nierengarten, B. Delavaux-Nicot, F. Monti, C. Chiorboli, N. Armaroli, *New J. Chem.* **2011**, *35*, 2234–2244.
- [84] U. Hahn, J.-F. Nierengarten, F. Vögtle, A. Listorti, F. Monti, N. Armaroli, *New J. Chem.* **2009**, *33*, 337–344.



- [85] M. Urbani, J. Iehl, I. Osinska, R. Louis, M. Holler, J.-F. Nierengarten, *Eur. J. Org. Chem.* **2009**, 2009, 3715–3725.
- [86] A. Bolag, H. Hayashi, P. Charbonnaz, N. Sakai, S. Matile, *ChemistryOpen* **2013**, 2, 55–57.
- [87] I. Nierengarten, J.-F. Nierengarten, *Chem. Rec.* **2015**, 15, 31–51.
- [88] M. Prato, Q. C. Li, F. Wudl, V. Lucchini, *J. Am. Chem. Soc.* **1993**, 115, 1148–1150.
- [89] Y. N. Yamakoshi, T. Yagami, S. Sueyoshi, N. Miyata, *J. Org. Chem.* **1996**, 61, 7236–7237.
- [90] J. Iehl, R. P. de Freitas, J.-F. Nierengarten, *Tetrahedron Lett.* **2008**, 49, 4063–4066.
- [91] T. T. Nguyen, S. Albert, T. L. A. Nguyen, R. Deschenaux, *RSC Adv.* **2015**, 5, 27224–27228.
- [92] J.-C. Wu, D.-X. Wang, Z.-T. Huang, M.-X. Wang, *J. Org. Chem.* **2010**, 75, 8604–8614.
- [93] R. Pereira de Freitas, J. Iehl, B. Delavaux-Nicot, J.-F. Nierengarten, *Tetrahedron* **2008**, 64, 11409–11419.
- [94] J. Iehl, I. Osinska, R. Louis, M. Holler, J.-F. Nierengarten, *Tetrahedron Lett.* **2009**, 50, 2245–2248.
- [95] G. R. Pereira, L. J. Santos, I. Luduvico, R. B. Alves, R. P. de Freitas, *Tetrahedron Lett.* **2010**, 51, 1022–1025.
- [96] A. Gmehling, A. Hirsch, *Eur. J. Org. Chem.* **2013**, 2013, 5093–5105.
- [97] J. Iehl, J.-F. Nierengarten, *Chem. – Eur. J.* **2009**, 15, 7306–7309.
- [98] P. Pierrat, S. Vanderheiden, T. Muller, S. Bräse, *Chem. Commun.* **2009**, 1748–1750.
- [99] J. Iehl, J.-F. Nierengarten, A. Harriman, T. Bura, R. Ziessel, *J. Am. Chem. Soc.* **2012**, 134, 988–998.
- [100] K. Yoosaf, J. Iehl, I. Nierengarten, M. Hmadeh, A.-M. Albrecht-Gary, J.-F. Nierengarten, N. Armaroli, *Chem. – Eur. J.* **2014**, 20, 223–231.
- [101] J. Iehl, M. Frasconi, H.-P. J. de Rouville, N. Renaud, S. M. Dyar, N. L. Strutt, R. Carmieli, M. R. Wasielewski, M. A. Ratner, J.-F. Nierengarten, et al., *Chem. Sci.* **2013**, 4, 1462–1469.
- [102] P. Fortgang, E. Maisonhaute, C. Amatore, B. Delavaux-Nicot, J. Iehl, J.-F. Nierengarten, *Angew. Chem. Int. Ed.* **2011**, 50, 2364–2367.
- [103] S. Guerra, J. Iehl, M. Holler, M. Peterca, D. A. Wilson, B. E. Partridge, S. Zhang, R. Deschenaux, J.-F. Nierengarten, V. Percec, *Chem. Sci.* **2015**, 6, 3393–3401.
- [104] D. Sigwalt, M. Holler, J. Iehl, J.-F. Nierengarten, M. Nothisen, E. Morin, J.-S. Remy, *Chem. Commun.* **2011**, 47, 4640–4642.
- [105] J.-F. Nierengarten, J. Iehl, V. Oerthel, M. Holler, B. M. Illescas, A. Muñoz, N. Martín, J. Rojo, M. Sánchez-Navarro, S. Cecioni, et al., *Chem. Commun.* **2010**, 46, 3860–3862.
- [106] P. Compain, C. Decroocq, J. Iehl, M. Holler, D. Hazelard, T. Mena Barragán, C. Ortiz Mellet, J.-F. Nierengarten, *Angew. Chem. Int. Ed.* **2010**, 49, 5753–5756.
- [107] M. Sánchez-Navarro, A. Muñoz, B. M. Illescas, J. Rojo, N. Martín, *Chem. – Eur. J.* **2011**, 17, 766–769.
- [108] J. Luczkowiak, A. Muñoz, M. Sánchez-Navarro, R. Ribeiro-Viana, A. Ginieis, B. M. Illescas, N. Martín, R. Delgado, J. Rojo, *Biomacromolecules* **2013**, 14, 431–437.
- [109] K. Buffet, E. Gillon, M. Holler, J.-F. Nierengarten, A. Imberty, S. P. Vincent, *Org. Biomol. Chem.* **2015**, 13, 6482–6492.

[110] I. Nierengarten, J.-F. Nierengarten, *Chem. – Asian J.* **2014**, 9, 1436–1444.



# **Chapter II: Regioselective syntheses of C<sub>60</sub> derivatives by a macrocyclic approach.**

## 1. Introduction.

The fullerene chemistry is now well understood and a large variety of reactions have been developed to functionalize fullerenes. The 30 double bonds of  $C_{60}$  being all equivalents, only one mono-adduct can be formed. In the case of further functionalization, the regiochemistry and the stereochemistry of the formed multi-adducts must be considered.

Problems related to the regiochemistry (achiral and chiral addition patterns) and the stereochemistry (in/out isomerism) considerably complicate the preparation of a specific multi-adducts of  $C_{60}$ . However with the specific reactivity of the *e* positions,  $T_h$ -hexa-adducts of  $C_{60}$  bearing twelve identical functional groups are accessible. Some examples of mixed hexa-adducts of  $C_{60}$  have been also reported (see **Chapter IV.1**), but the lack of post-functionalization possibilities, low yields or fastidious syntheses still limits the preparation of highly functionalized mixed hexa-adducts. The straightforward syntheses of mixed hexa-adducts pass through the elaboration of a simple regio- and stereoselective methodology for the formation of bis- or tris-adducts of  $C_{60}$  with appropriate addition patterns.

Two main approaches have been developed for the regioselective functionalization of  $C_{60}$ , the tether-directed method which will be discussed in **Chapter III** and the macrocyclic-directed method. In this last approach, the malonates are included within a macrocyclic structure (Figure II-1). The conformation and the size of the macrocycle will then play an important role in the regioselectivity of the cyclization on  $C_{60}$ .

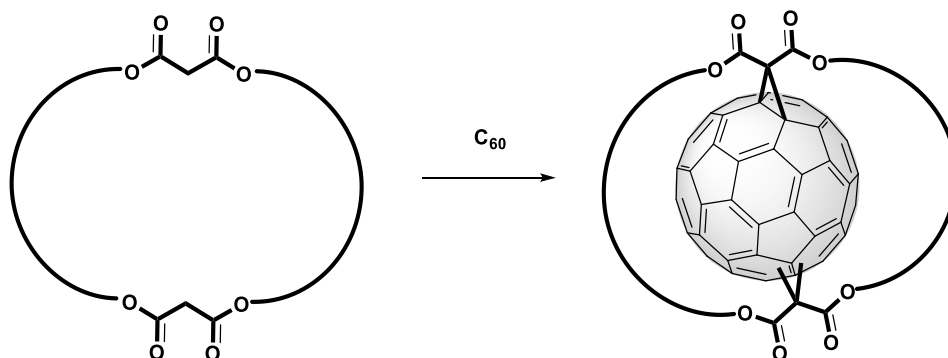


Figure II-1 : Macrocyclic-directed method for the region-functionalization of  $C_{60}$ .

The macrocyclic-directed method was introduced in 2002 by Hirsch and coworkers.<sup>[1]</sup> With this method, they were able to obtain high regioselectivity and good yields for the functionalization of  $C_{60}$ . The length of the spacer and more generally the ring size of the macrocycle are determinant to obtain a specific isomer. Indeed for bis-adducts of  $C_{60}$ , no regioselectivity was observed for a 26-membered ring macrocyclic bis-malonate. In contrast, by increasing the size of the macrocycle, they were able to obtain the *trans*-3 and *trans*-1 bis-adducts of  $C_{60}$  in a regioselective manner. For example, cyclo-[2]-dodecylmalonate gave exclusively the *trans*-3 bis-adducts with a yield of 56% (Figure II-2). They also synthesized mixed macrocycles with two alkyl chains of different length. With unequal distances between

the ester functions, only bis-adducts of  $C_{60}$  with  $C_5$ -symmetrical addition pattern were obtained. For example, cyclo-[2]-octyl-tetradecylmalonate gave exclusively the *e* bis-adducts in 51% yield.

An increased number of units in the macrocycle play also an important role on the regioselectivity by an enlargement of the ring size. No regioselectivity was observed for the cyclo-[2]-octylmalonate but in contrast, when the reaction was carried out with the cyclo-[3]-octylmalonate, tris-adducts *e,e,e* and *trans-4,trans-4,trans-4* were obtained with 40% and 2% yields, respectively (Figure II-2). Another tris-adducts (*trans-3,trans-3,trans-3*) was obtained in 30% yield with a cyclo-[3]-tetradecylmalonate.

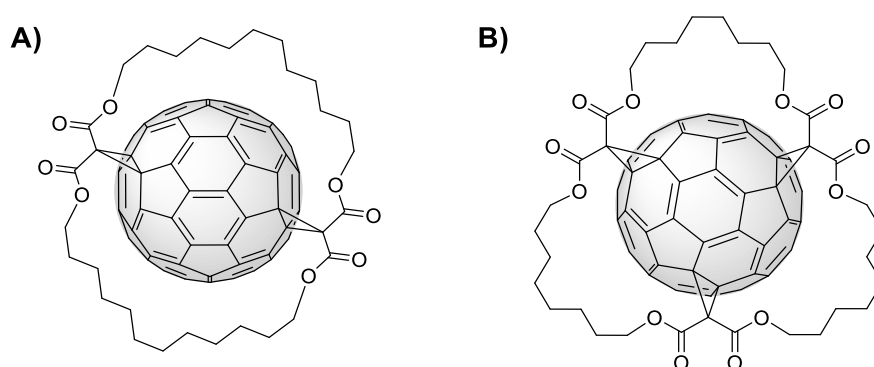


Figure II-2 : Examples of bis- and tris-adducts obtained by a macrocyclic approach. A) Bis-adducts *trans-3*. B) Tris-adducts *e,e,e*.

Wilson and coworkers reported syntheses using the same methodology, but instead of alkyl chains, they used oligoglycols as flexible linkers.<sup>[2]</sup> High regioselectivity and good yields were still obtained. Bis- or tris-adducts with rotational symmetry are favored when symmetric macrocycles are used. Nonetheless, a bis-adducts *e* ( $C_5$ -symmetrical addition pattern) (Figure II-3) was also obtained in addition to a *cis-3* bis-adducts with tetraethylene glycol linkers. They also reported an original *trans-4,cis-3,cis-3* tris-adducts that was isolated with a yield of 51%.

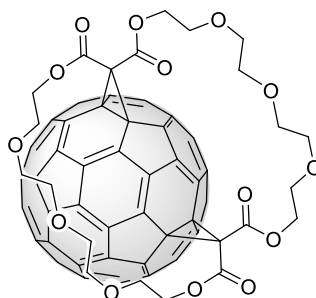


Figure II-3 : Example of a  $C_5$ -symmetrical addition pattern of a bis-adducts *e*.

Some addition patterns are inherently chiral (bis-adducts *cis-3*, *trans-3* and *trans-2* or tris-adducts *e,e,e*) and these fullerene multi-adducts are obtained as racemic mixtures. Purification of enantiomerically pure  $C_{60}$  derivatives requires fastidious separations by preparative chromatography (HPLC) on chiral stationary phases.<sup>[3]</sup>

Chronakis and Hirsch achieved the separation of enantiomerically pure  $^fC$  and  $^fA$  addition patterns of *trans*-3 and *e,e,e* regioisomers (Figure II-4).<sup>[4,5]</sup> Starting from optically pure spacers, they synthesized the corresponding chiral cyclo-oligo-malonates. Reaction of the optically pure cyclo-oligo-malonates with  $C_{60}$  afforded diastereoisomeric mixtures of bis- or tris-adducts according to the used macrocycle. The difference of polarity of the diastereoisomers made possible the separation of enantiomerically pure *trans*-3 and *e,e,e* addition pattern by classical silica column chromatography. Diastereoselectivity was also observed between the  $^fC$  and  $^fA$  addition pattern.

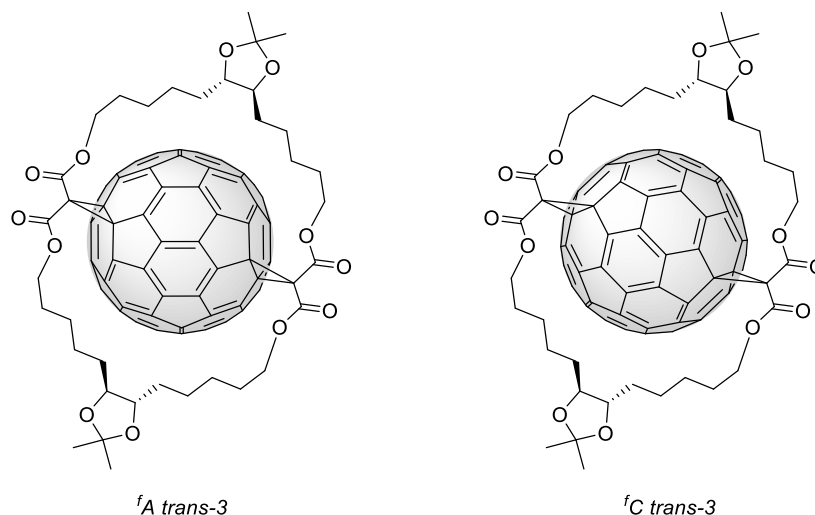


Figure II-4 : Example of inherently chiral addition pattern of  $C_{60}$  derivatives.

In the last example, Nierengarten and coworkers have shown that a rigid macrocycle is also suitable for the regioselective functionalization of  $C_{60}$ .<sup>[6]</sup> By the means of a phenylethynyl spacer and a silyl group, a rigid macrocyclic bis-malonate was synthesized. Subsequent reaction with  $C_{60}$  afforded the *trans*-3 and *trans*-1 bis-adducts in a 2:1 ratio (Figure II-5). Unfortunately, it was impossible to remove the silyl groups in this case in order to further functionalize these fullerene bis-adducts.

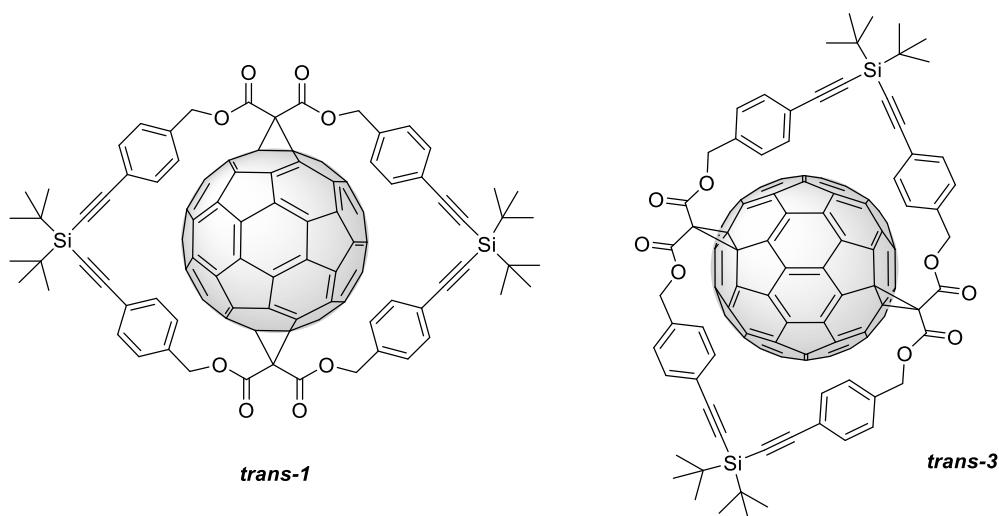


Figure II-5 : Bis-adducts obtained with a rigid macrocycle.

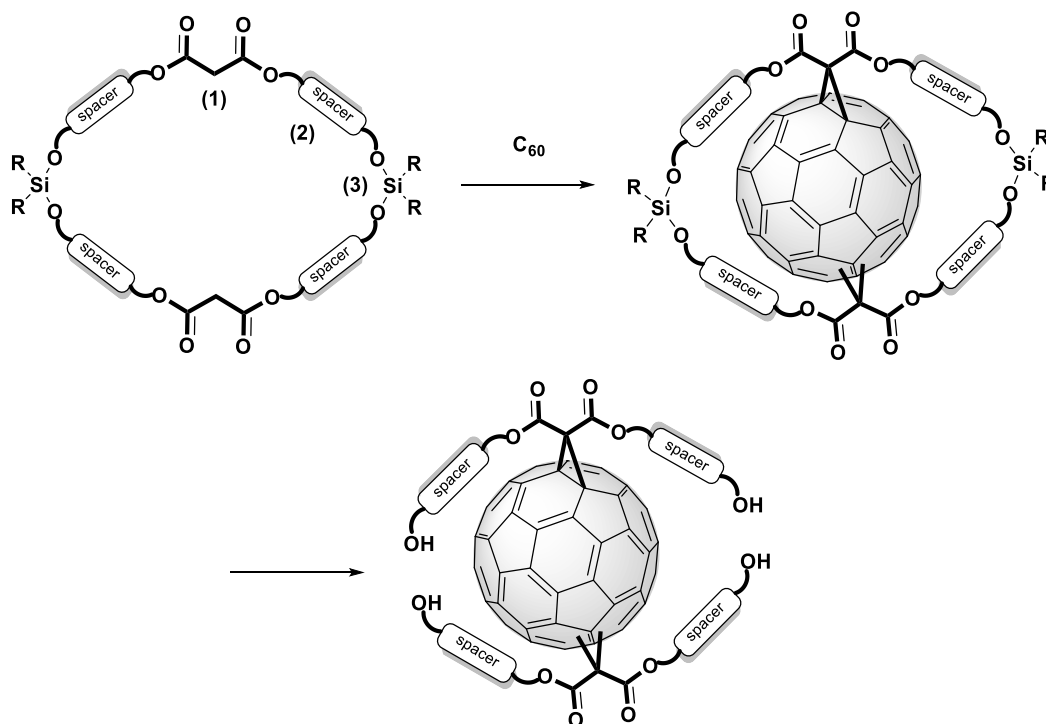


Figure II-6 : Concept of macrocycle incorporating silyl groups.

The macrocyclic approach affords good results for the regioselective functionalization of  $C_{60}$ . However, the post-functionalization of the addends bond on the  $C_{60}$  is difficult. In this context, the introduction of easily removable sub-units in the macrocycle is needed for an easy post-functionalization of these fullerene derivatives. Preliminary results obtained in our laboratory have shown that silyl groups as alcohol protecting group can be easily removed. In this work, the use of silyl groups as alcohol protecting group within a macrocycle is presented for the regioselective functionalization of  $C_{60}$ . The conception of the macrocycle will be based on three elements (Figure II-6): (1) malonates for anchoring on the  $C_{60}$ ; (2) diol spacers to reach specific positions on the  $C_{60}$ ; (3) removable silyl groups to allow the post-functionalization of the multi-adducts of  $C_{60}$ . In order to study the regioselectivity of macrocyclic bis-malonates incorporating silyl groups, three parameters have been modified: (1) the ring size; (2) the silyl group and (3) the flexibility of the spacer.



## 2. Regioselective syntheses of Bis-adducts of C<sub>60</sub> by macrocyclic approach.

### 2.1 Preparation of macrocyclic bis-malonates.

#### 2.1.1 Synthesis of macrocyclic bis-malonates: the direct approach.

The first challenge has been to develop an efficient synthetic route for the macrocyclic oligo-malonates incorporating silane subunits. A first approach was based on the direct macrocyclization of diols with malonyl chloride (Figure II-7). The diols are either obtained by reduction of the corresponding diesters or by reaction of a monoprotected diol with an appropriated silylating agent followed by deprotection.

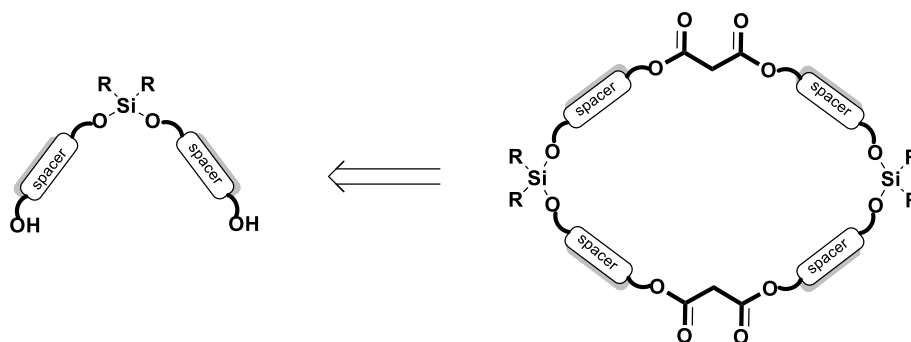
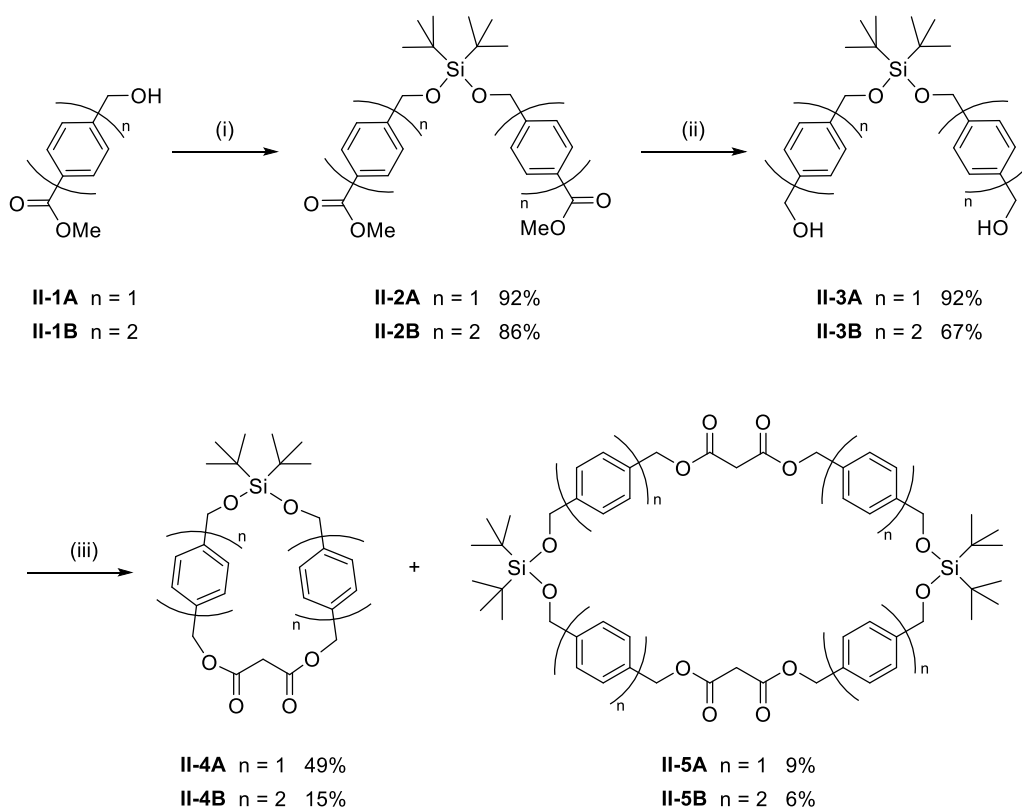


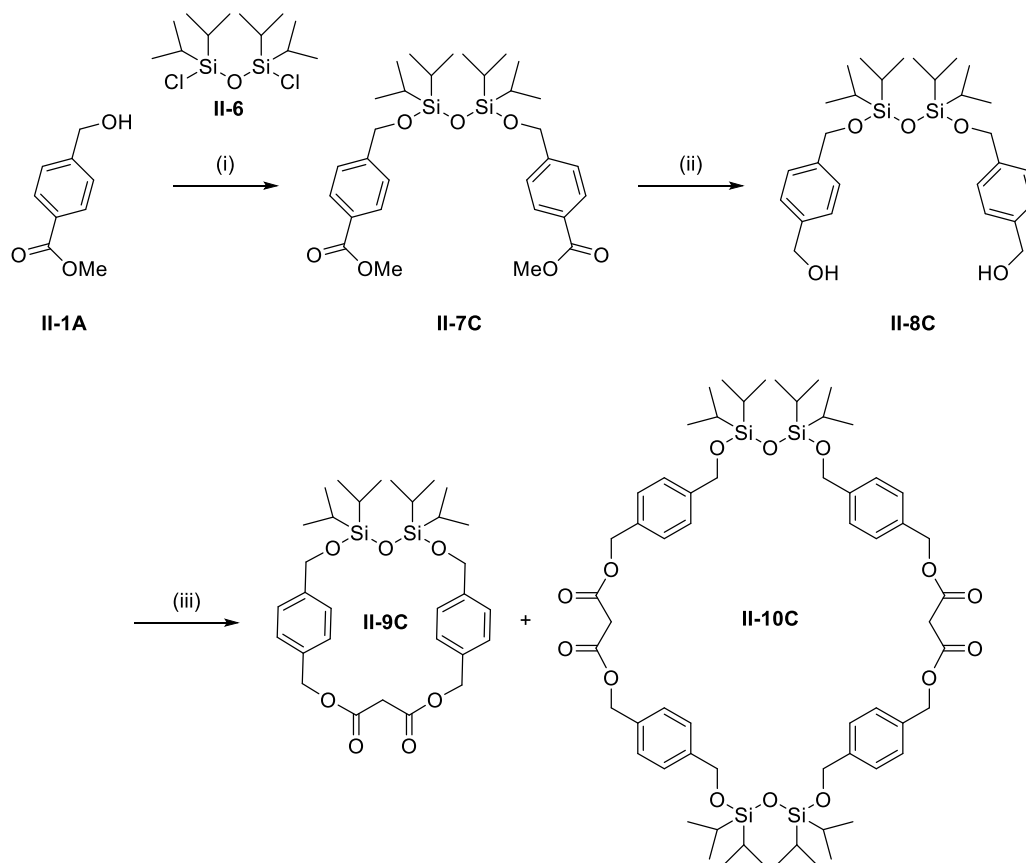
Figure II-7 : Retrosynthetic analysis for the elaboration of cyclo-bis-malonate

The preparation of a first macrocyclic bis-malonate with rigid spacers (**II-5**) is depicted in **Scheme II-1**. Treatment of **II-1** with di-*t*-butylsilylene bis(trifluoromethanesulfonate) ( $t\text{Bu}_2\text{Si}(\text{OTf})_2$ ) in DMF in the presence of pyridine gave compounds **II-2**. The reduction of the ester functions of **II-2A** was first attempted with  $\text{LiAlH}_4$ . Under these conditions, the desired diol (**II-3A**) was not obtained due to the cleavage of the silyl ethers. In contrast, when diisobutylaluminium hydride (DIBAL-H) was used as the reducing agent, compound **II-3A** was obtained in good yield (92%). Similarly, treatment of **II-2B** with DIBAL-H in  $\text{CH}_2\text{Cl}_2$  at  $-15^\circ\text{C}$  afforded diol **II-3B** in 67% yield. Subsequent reactions of diols **II-3** with malonyl chloride were performed in the presence of 4-dimethylaminopyridine (DMAP) under pseudo high dilution conditions. Based on the order of elution,<sup>[1]</sup> cyclomonomalones **II-4** and cyclobismalonates **II-5** were isolated.



**Scheme II-1.** Reagents and conditions: (i)  $t\text{Bu}_2\text{Si}(\text{OTf})_2$ , DMF, pyridine, rt, 12 h; (ii) DIBAL-H,  $\text{CH}_2\text{Cl}_2$ ,  $-15^\circ\text{C}$ , 3 h; (iii) malonyl chloride, DMAP,  $\text{CH}_2\text{Cl}_2$ , rt, 2 h.

To study the influence of the silyl group, the macrocyclic bis-malonates **II-10C** was synthesized from the silyl group 1,3-dichloro-1,1,3,3-tetraisopropyldisiloxane (**II-6**). Compound **II-10C** was prepared by following the synthetic route developed for the preparation of compounds **II-5** and was obtained in 5% yield (**Scheme II-2**).



**Scheme II-2.** Reagents and conditions: (i) **II-6**, DMF, pyridine, rt, 12 h (99%); (ii) DIBAL-H, CH<sub>2</sub>Cl<sub>2</sub>, -15°C, 3 h (77%); (iii) malonyl chloride, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, rt, 2 h (**II-9C**: 73%; **II-10C**: 5%).

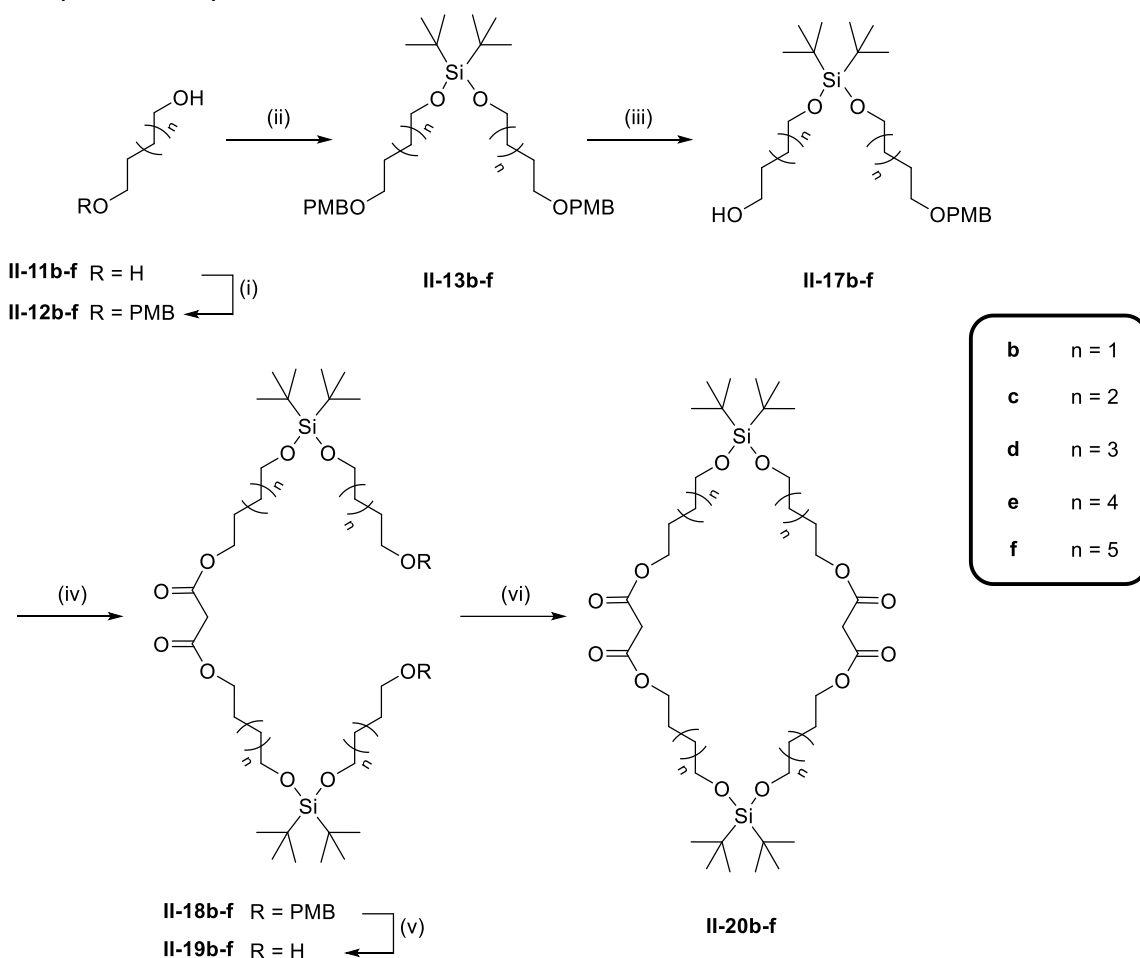
Macrocycles with alkyl spacers have been also prepared. The synthesis of the macrocyclic bis-malonate **II-16a** shown in **Scheme II-3** was first achieved. The selective mono-protection of 1,3-propanediol was carried out by treatment with Ag<sub>2</sub>O and *p*-methoxybenzyl chloride (PMBCl) according to the conditions reported by Bouzide and Sauv e.<sup>[7]</sup> Treatment of the resulting mono-protected derivative **II-12a** with *t*-Bu<sub>2</sub>Si(OTf)<sub>2</sub> and imidazole in DMF provided **II-13a** in 68% yield. The PMB protecting groups in **II-13a** were conveniently removed by treatment with 2,3-dichloro-5,6-dicyanobenzoquinone (DDQ) in CH<sub>2</sub>Cl<sub>2</sub> containing a small amount of water. The di-*t*-butylsilylene group remained unchanged and key building block **II-14a** was thus obtained in good yields (86%). Reaction of diol **II-14a** with malonyl chloride in the presence of DMAP under pseudo high dilution conditions afforded a mixture of cyclooligomers and polymers. Macrocycles **II-15a** and **II-16a** were isolated in a pure form in 24% and 15% yield, respectively. In the case of **II-16a**, the proposed structure was confirmed by MALDI-TOF mass spectrometry.



## 2.1.2 Synthesis of macrocyclic bis-malonates: the stepwise approach.

This work has been done in collaboration with Dr. Thi Minh Nguyet Trinh and Dr. Sebastiano Guerra.

Larger macrocyclic oligo-malonates were prepared with alkanediols with 4 to 8 carbon atoms. The stepwise synthetic route for the preparation of these macrocyclic bis-malonates **II-20** is depicted in **Scheme II-5**. Mono-protection of diols **II-11b-f** by treatment with PMBCl/Ag<sub>2</sub>O followed by silylation (*t*Bu<sub>2</sub>Si(OTf)<sub>2</sub>/imidazole) afforded compound **II-13b-f**. Treatment of **II-13** with 1 equiv. of DDQ provided the mono-protected derivatives **II-17**. Subsequent reaction with malonyl chloride in the presence of DMAP gave malonates **II-18**. The PMB protecting groups being removed under neutral conditions by treatment with DDQ, the silyl ether functions remained effectively intact and diols **II-19** were obtained in good yields (68 to 85%). Reaction of **II-19** with malonyl chloride under pseudo high dilution conditions gave cyclobismalonates **II-20** in 19 to 50% yields. Macrocycles **II-20**, which have ring sizes of 32-36-40-44-48 atoms respectively, were characterized by NMR spectroscopy and mass spectrometry.



**Scheme II-5.** Reagents and conditions: (i) PMBCl, Ag<sub>2</sub>O, CH<sub>2</sub>Cl<sub>2</sub>, 48 h (69-95%); (ii) *t*Bu<sub>2</sub>Si(OTf)<sub>2</sub>, DMF, imidazole, rt, 12 h (60-91%); (iii) DDQ (1 eq), CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, rt, 6 h (47-50%); (iv) malonyl chloride, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, rt, 2 h (67-96%); (v) DDQ (2.5 eq), CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, rt, 2 h (68-85%); (vi) malonyl chloride, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, rt, 2 h (19-50%).

## 2.2 Regioselective functionalization of C<sub>60</sub> with macrocyclic bis-malonates.

Reaction of macrocyclic bis-malonates with C<sub>60</sub> were carried out under the typical conditions developed by Diederich *et. al.* for the preparation of fullerene bis-adducts (**Scheme II-6**).<sup>[8]</sup> Specifically, 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU) was added to a solution of cyclo-bis-malonates, C<sub>60</sub> and iodine in toluene at room temperature.

Reaction of **II-5A** with C<sub>60</sub> was highly regioselective and only the *trans*-3 regioisomer was obtained in a good yield of 54%. In contrast, no regioselectivity was observed for the larger macrocycle **II-5B**. Bis-adducts *trans*-3, *e* and *cis*-2 were isolated in 15%, 4% and 6% yields, respectively. Traces of other bis-adducts were also observed but too small amounts of products prevented complete characterizations. The Bingel reaction with macrocycle **II-10C** was regioselective and afforded the *trans*-3 (**II-21**) and the *e* (**II-22**) bis-adducts in 26% and 15% yields, respectively.

Reaction of the 28 and 32-membered ring macrocyclic bis-malonates (**II-16a** and **II-20b**) with C<sub>60</sub> were also highly regioselective and gave the *cis*-2 C<sub>60</sub> bis-adduct **II-23** in 27% and 38% yield, respectively. A further increase of the ring size with the 36-48 membered ring macrocyclic bis-malonates **II-20c-f** afforded the two regioisomeric bis-adducts *trans*-3 (**II-21c-f**) and *e* (**II-22c-f**) by reaction with C<sub>60</sub>.

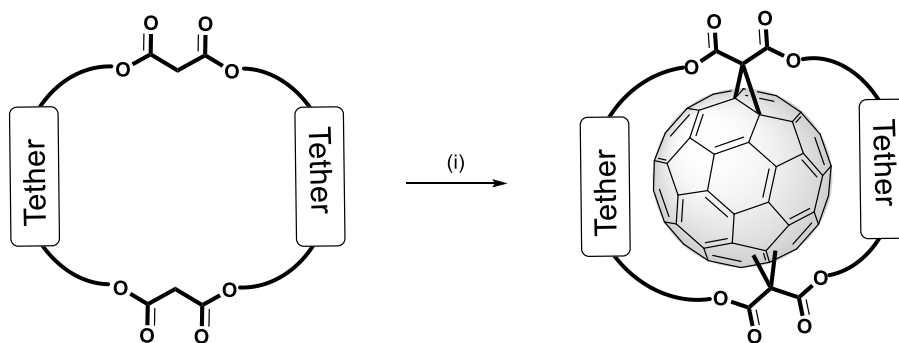
Compounds **II-21-23** have been characterized by <sup>1</sup>H and <sup>13</sup>C NMR, UV-vis and IR spectroscopies. In addition, the MALDI-TOF mass spectra were also consistent with the proposed structures. The relative position of the two cyclopropane rings on the fullerene core were determined based on the molecular symmetry deduced from the NMR spectra and on the diagnostic features seen in the absorption spectrum.

In the case of a bis-functionalization of C<sub>60</sub> with macrocycles, there are four possible molecular symmetries based on the addition patterns: C<sub>1</sub> (*e*), C<sub>s</sub> (*cis*-2 and *trans*-4), C<sub>2</sub> (*cis*-3, *trans*-3 and *trans*-2) and D<sub>2h</sub> (*trans*-1). To deduce the molecular symmetry, careful analyses of the fullerene resonances on the <sup>13</sup>C NMR spectra of **II-21-23** were performed. As typical examples, the <sup>13</sup>C NMR spectra of an *e*, a *trans*-3 and a *cis*-2 regioisomers are shown in [Figure II-8](#).

In the case of **II-22c**, 60 fullerene resonances were observed: four are observed at  $\delta = 69.7, 71.35, 71.37$  and  $71.4$  ppm (sp<sup>3</sup> C atoms) and 56 between  $138.9$  and  $148.1$  ppm (sp<sup>2</sup> C atoms) indicating a C<sub>1</sub> symmetry.

In the case of the *trans*-3 bis-adducts **II-21**, 30 fullerene resonances are expected. Indeed, for compound **II-21c** two are observed at  $\delta = 71.0$  and  $70.9$  ppm (sp<sup>3</sup> C atoms) and 28 between  $\delta = 139.4$  and  $146.7$  ppm (sp<sup>2</sup> C atoms) which is in perfect agreement with a C<sub>2</sub> molecular symmetry.

For compound **II-23a**, two fullerene resonances are observed at  $\delta = 67.6$  and  $70.5$  ppm (sp<sup>3</sup> C atoms) and 30 between  $\delta = 135.7$  and  $148.2$  ppm (sp<sup>2</sup> C atoms). The presence of half intensity signals for some of the resonances of the fullerene C atoms seen in the sp<sup>2</sup> region is an unambiguous signature for a C<sub>s</sub> symmetrical structure.



	Tether	Regioisomer of C <sub>60</sub> bis-adduct	Yield
II-5A		<i>trans</i> -3	II-21A 54%
II-5B		<i>trans</i> -3 <i>e</i> <i>cis</i> -2	II-21B 15% II-22B 4% II-23B 6%
II-10C		<i>trans</i> -3 <i>e</i>	II-21C 26% II-22C 15%
II-16a		<i>cis</i> -2	II-23a 27%
II-20b		<i>cis</i> -2	II-23b 38%
II-20c		<i>trans</i> -3 <i>e</i>	II-21c 44% II-22c 12%
II-20d		<i>trans</i> -3 <i>e</i>	II-21d 33% II-22d 27%
II-20e		<i>trans</i> -3 <i>e</i>	II-21e 30% II-22e 20%
II-20f		<i>e</i> <i>trans</i> -3	II-22f 21% II-21f 16%

Scheme II-6. Reagents and conditions: (i) C<sub>60</sub>, I<sub>2</sub> and DBU, PhMe, rt, 1 h.

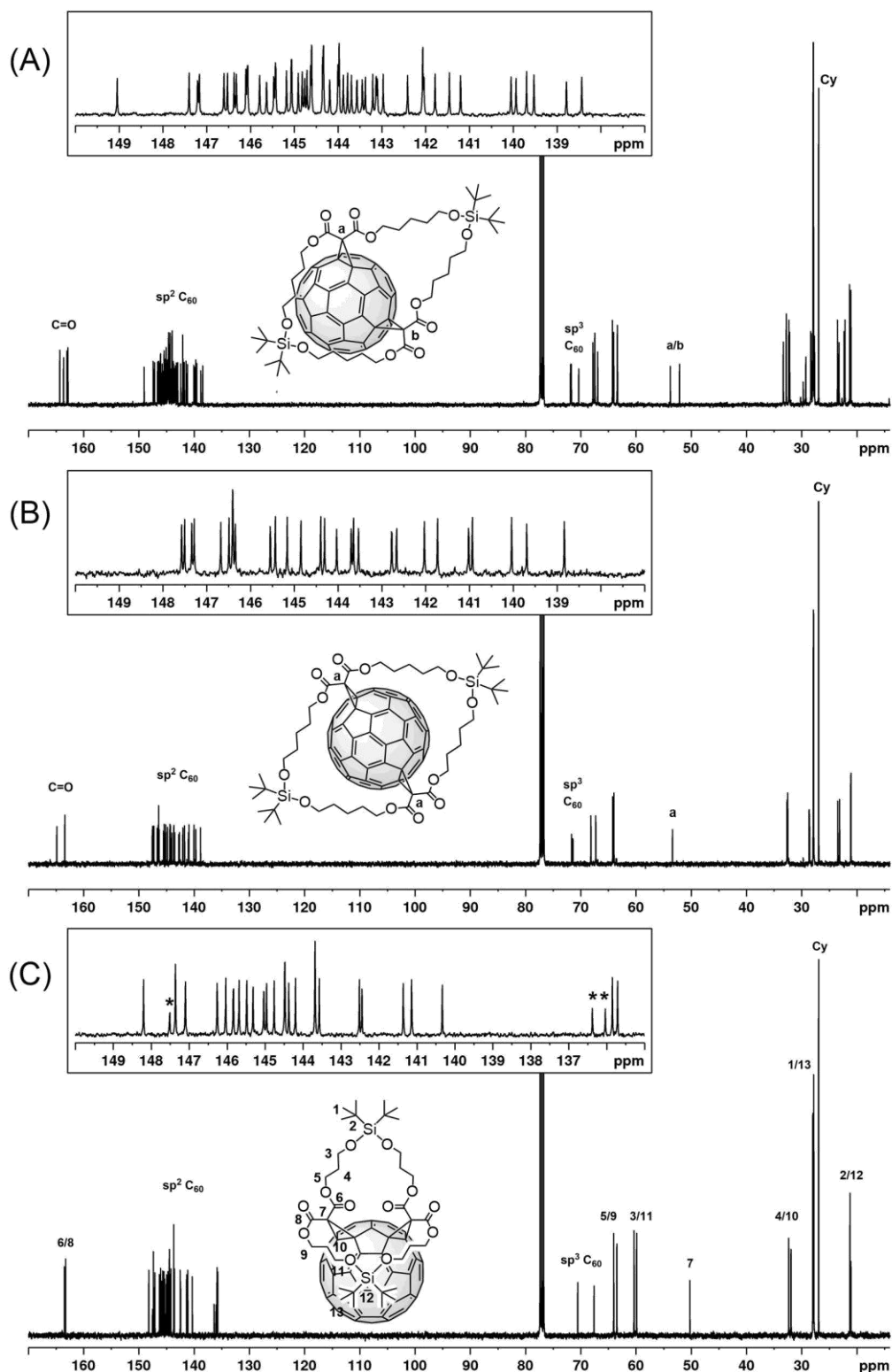


Figure II-8:  $^{13}\text{C}$  NMR spectra ( $\text{CDCl}_3$ , 100 MHz); (A)  $C_1$ -symmetrical *e* bis-adduct **II-22c**; (B)  $C_2$ -symmetrical *trans*-3 bis-adduct **II-21c**; (C)  $C_5$ -symmetrical *cis*-2 bis-adduct **II-23a**; Inset: detailed view showing the resonances of the fullerene  $\text{sp}^2 \text{C}$  atoms; \* indicates the  $\text{sp}^2$  fullerene C atoms showing half intensity signals; Cy = cyclohexane.



The unambiguous assignation of the addition patterns was finally achieved by UV-vis analysis. The absorption spectra of C<sub>60</sub> multi-adducts are highly dependent on the addition pattern and characteristic for each of the regioisomers (Figure II-9).<sup>[8-13]</sup> The absorption spectra of **II-21-23** were recorded in CH<sub>2</sub>Cl<sub>2</sub> and clearly revealed the diagnostic features of *trans*-3, *e* and *cis*-2 addition patterns, respectively.

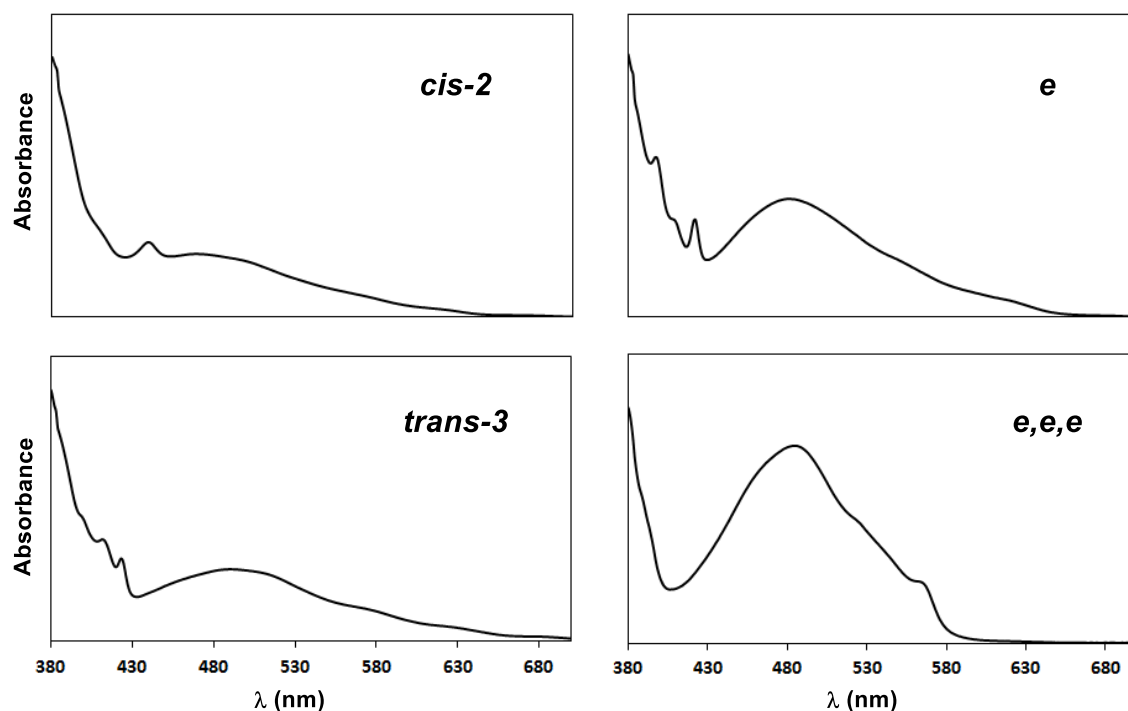


Figure II-9: Characteristic UV/vis spectra of four multi-adducts of C<sub>60</sub>.

The regioselectivity of the bis-addition on the C<sub>60</sub> is clearly governed by the ring size of the macrocycle. Indeed, an increase of the macrocycle size induces a decrease in the regioselectivity.

The silyl group has also an impact on the regioselectivity. The fact that three different regioisomers were isolated with the macrocycle **II-5B** shows that the O-Si-O sequence introduces too much flexibility into the macrocycle. Indeed, the macrocycle must adopt different conformations in order to form these regioisomers. To adopt these different conformations, the necessary degrees of freedom are provided by the silyl groups. This is further validated by the formation of the *e* and *trans*-3 regioisomers with macrocycle **II-10C**.

The influence of the rigidity/flexibility of the spacer on the regioselectivity is only minor to the fact that the silyl groups provide flexibility to the macrocycle.

The preferential formation of the *e* and *trans*-3 C<sub>60</sub> bis-adducts is explained by the reactivity of the C<sub>60</sub> sphere. For C<sub>60</sub> mono-adduct derivatives, the *e* and *trans*-3 positions are the most reactive sites.<sup>[9,14]</sup> Therefore bis-adducts *e* and *trans*-3 are kinetic products. In counterpart, the thermodynamic product is formed when the formation of *e* and *trans*-3 bis-adducts are energetically disfavored by strain arguments relative to the conformation of the

macrocycle. In the case of the cyclo-alkylmalonates, computational calculations at the AM1 semi-empirical level were performed with Spartan'10 Macintosh Parallel Edition (Wavefunction Inc., USA) (Table II-1). As shown in Table II-1, the *e* and *trans*-3 isomers are low in energy for (CH<sub>2</sub>)<sub>5-8</sub> and confirmed the observed regioselectivity. The preference for the C<sub>2</sub>-symmetrical C<sub>60</sub> bis-adducts *trans*-3 can be explained by the higher molecular symmetry. In general, more a molecule is symmetric, more the strain is uniformly distributed and thus provides a more stable compound. In the case of (CH<sub>2</sub>)<sub>3-4</sub>, the formation of the *e* and *trans*-3 isomers are energetically disfavored compared to the *cis*-2 isomer and thus only the *cis*-2 bis-adduct of C<sub>60</sub> were formed.

	(CH <sub>2</sub> ) <sub>3</sub> II-16a	(CH <sub>2</sub> ) <sub>4</sub> II-20b	(CH <sub>2</sub> ) <sub>5</sub> II-20c	(CH <sub>2</sub> ) <sub>6</sub> II-20d	(CH <sub>2</sub> ) <sub>7</sub> II-20e	(CH <sub>2</sub> ) <sub>8</sub> II-20f
<i>cis</i> -2 C <sub>s</sub>	0	0	0	0	+3	+20.6
<i>cis</i> -3 C <sub>2</sub>	+180.3	+97.2	+121	+91.5	+79.8	+85.7
<i>e</i> C <sub>1</sub>	+76.3	+30.8	+18.1	+7.3	+1.1	+12.1
<i>trans</i> -4 C <sub>s</sub>	326.5	+57.2	+28.4	+16.2	+14.5	+23.1
<i>trans</i> -3 C <sub>2</sub>	+83.4	+30.3	+15.3	+21.3	0	+6.5
<i>trans</i> -2 C <sub>2</sub>		+219.2	+77.1	+53.2	+28	+36.8
<i>trans</i> -1 D <sub>2h</sub>			+71.3	+5.8	+39	+0

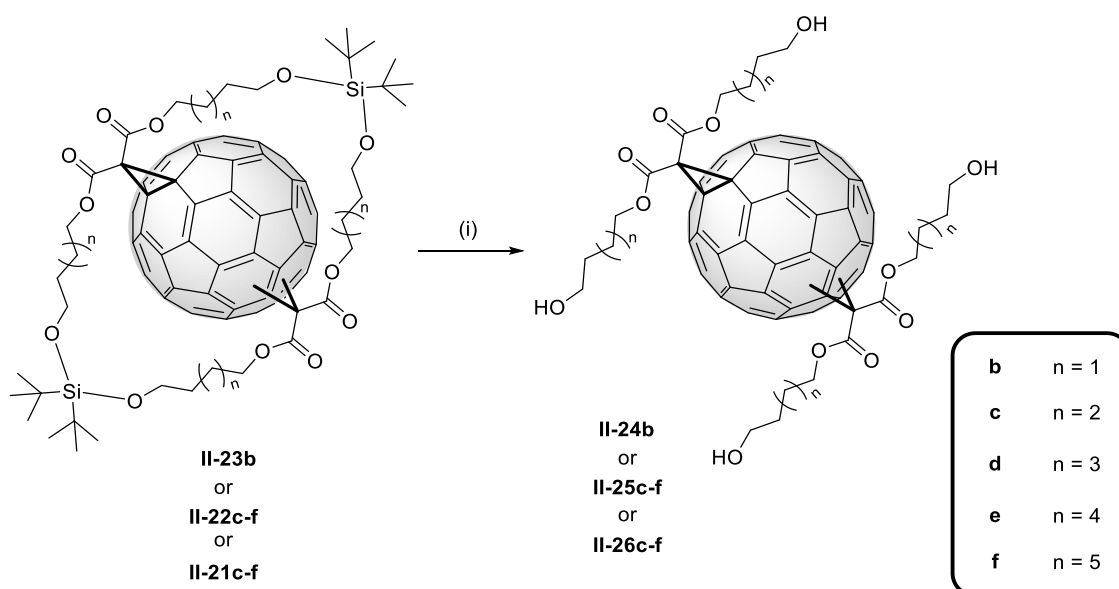
Table II-1 : Calculated  $\Delta E$  heat of formation in kJ/mol at the AM1 semi-empirical level. In green the major formed regioisomer and in orange the minor formed regioisomer.

### 2.3 Deprotection.

As seen previously, the silyl groups have an impact on the regioselectivity. However, the principal interest to integrate them into a macrocycle structure is their easy deprotection. In order to preserve the ester functions, basic deprotection conditions are excluded. The deprotection of compounds II-23b, II-22c-f and II-21c-f were carried out with an excess of BF<sub>3</sub>.Et<sub>2</sub>O in CH<sub>2</sub>Cl<sub>2</sub>/CH<sub>3</sub>CN (Scheme II-7). With acidic conditions, the ester functions remained unchanged and C<sub>60</sub> bis-adducts II-24, II-25 and II-26 were obtained in good yields (56-89%). The removal of the silyl groups was confirmed by the spectroscopic and spectrometric data. In the case of C<sub>1</sub>-symmetrical bis-adducts II-22, the removal of the silyl groups led to an increase of the molecular symmetry. A C<sub>s</sub>-symmetry was deduced from their <sup>13</sup>C NMR spectra.

As typical example, the <sup>13</sup>C NMR spectrum of **II-22c** is shown in [Figure II-10](#). Out of the 32 expected fullerene resonances, three are observed at  $\delta = 70.4, 71.58$  and  $71.65$  ppm for the sp<sup>3</sup> C atoms and 28 between  $\delta = 138.7$  and  $147.6$  ppm for the sp<sup>2</sup> C atoms. The presence of a half intensity signal in the sp<sup>2</sup> region is an unambiguous proof for the C<sub>5</sub>-symmetry of compound **II-22c**. Furthermore, the presence of three sp<sup>3</sup> fullerene C atoms resonances is an unambiguous confirmation of the *equatorial* addition pattern. The *e* C<sub>60</sub> bis-adduct is the sole C<sub>5</sub>-symmetrical regioisomer for which three sp<sup>3</sup> fullerene C atoms resonances are expected. Only two signals are expected for the two other possible C<sub>5</sub>-symmetrical regioisomers (*cis*-2 and *trans*-4).

Our macrocyclic bis-malonates approach have furnished three regioisomers with good regioselectivity. The silyl groups can be readily cleaved to afford the corresponding acyclic fullerene polyols. With their four alcohol functions, compound **II-24-26** are valuable building blocks for further chemical modifications. Bis-adducts **II-25** with its *e* addition pattern can also be used as precursor for the preparation of hexa-adducts of C<sub>60</sub> with an octahedral addition pattern.



**Scheme II-7.** Reagents and conditions: (i) BF<sub>3</sub>.Et<sub>2</sub>O, CH<sub>2</sub>Cl<sub>2</sub>, CH<sub>3</sub>CN, rt, 12 h (**II-24**: 88%; **II-25**: 70-85%; **II-26**: 56-89%)

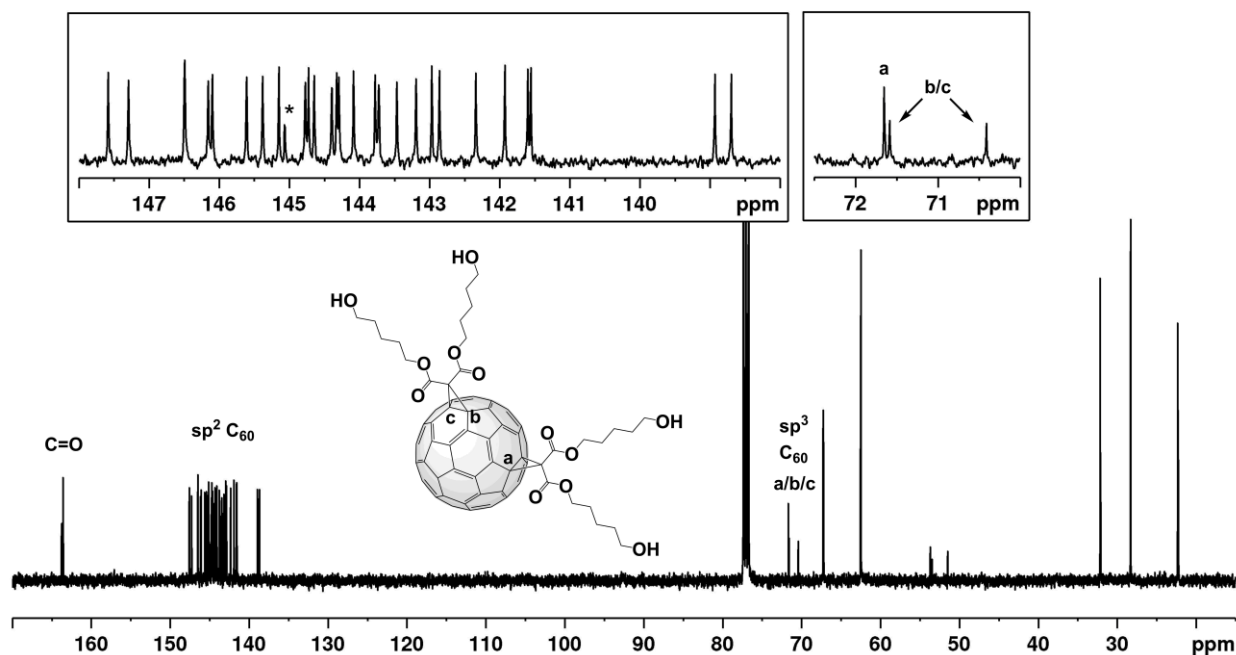


Figure II-10: <sup>13</sup>C NMR spectra (CDCl<sub>3</sub>, 100 MHz) of C<sub>5</sub>-symmetrical e bis-adduct **II-25c**; Inset: detailed view showing the resonances of the fullerene sp<sup>2</sup> C atoms; \* indicates the sp<sup>2</sup> fullerene C atoms showing half intensity signals.

### 3. Regioselective syntheses of Tris-adducts of C<sub>60</sub> by macrocyclic approach.

#### 3.1 Stepwise preparation of cyclo-tris-malonates.

This work has been done in collaboration with Eric Meischner.

Following the successful preparation of fullerene bis-adducts from macrocyclic bis-malonates, we became interested in extending our work to the synthesis of macrocyclic tris-malonates to prepare specific fullerene tris-adducts. For this purpose, an efficient synthetic route allowing the preparation of macrocyclic tris-malonates had to be developed first.

To selectively prepare cyclo-tris-malonates, we proposed to synthesize step-by-step the diol **II-33** containing two malonate sub-units (Figure II-11). The third malonate sub-unit being introduced by macrocyclisation reaction with malonyl chloride. In principle, diol **II-33** can be prepared from compounds **II-17** and **II-27** by an esterification reaction followed by deprotection. This last compounds are obtained from the protected derivative **II-13**. In a first instance, the synthesis of the cyclo-tris-malonate will be optimize with the (CH<sub>2</sub>)<sub>3</sub> spacer.

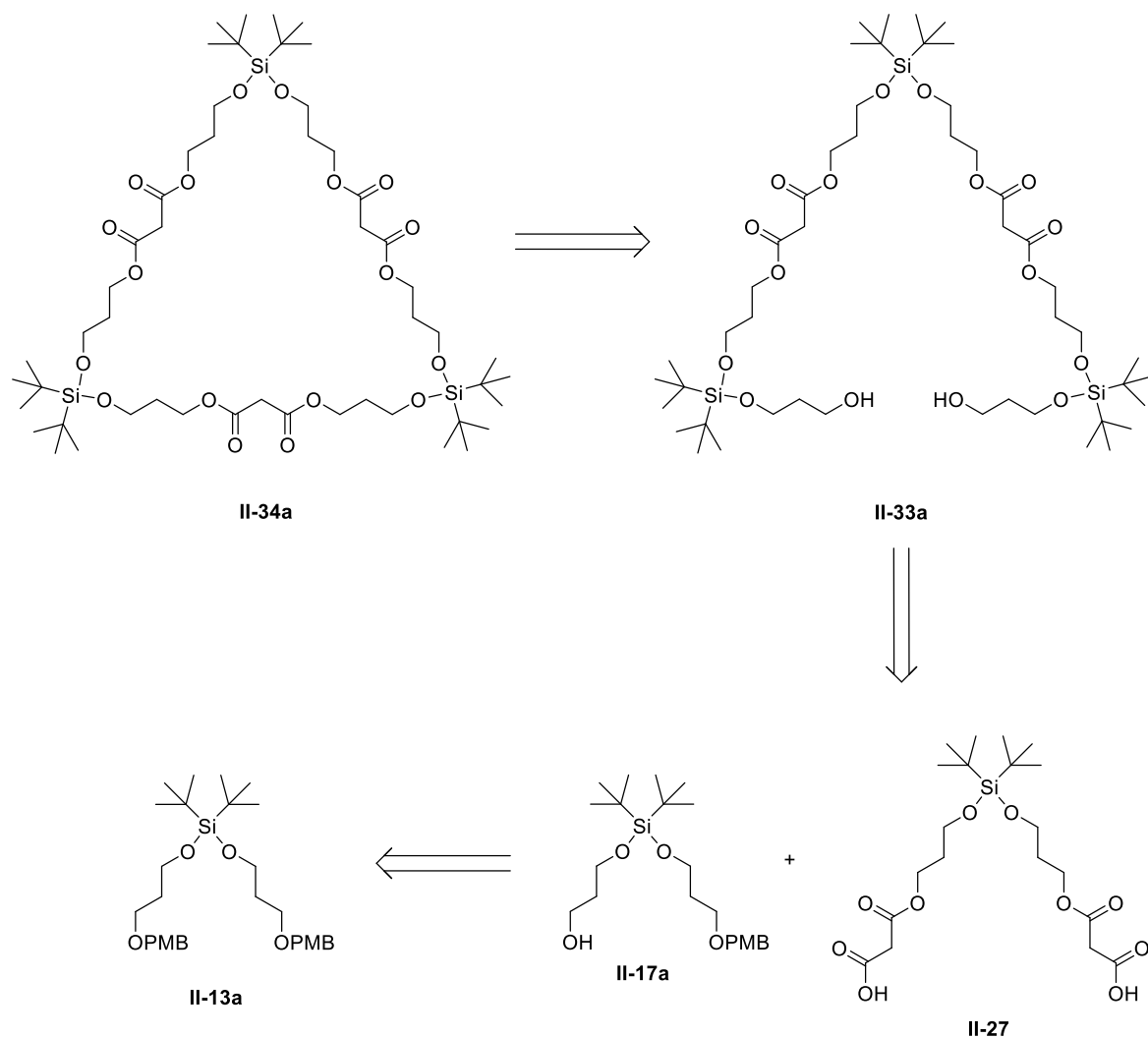
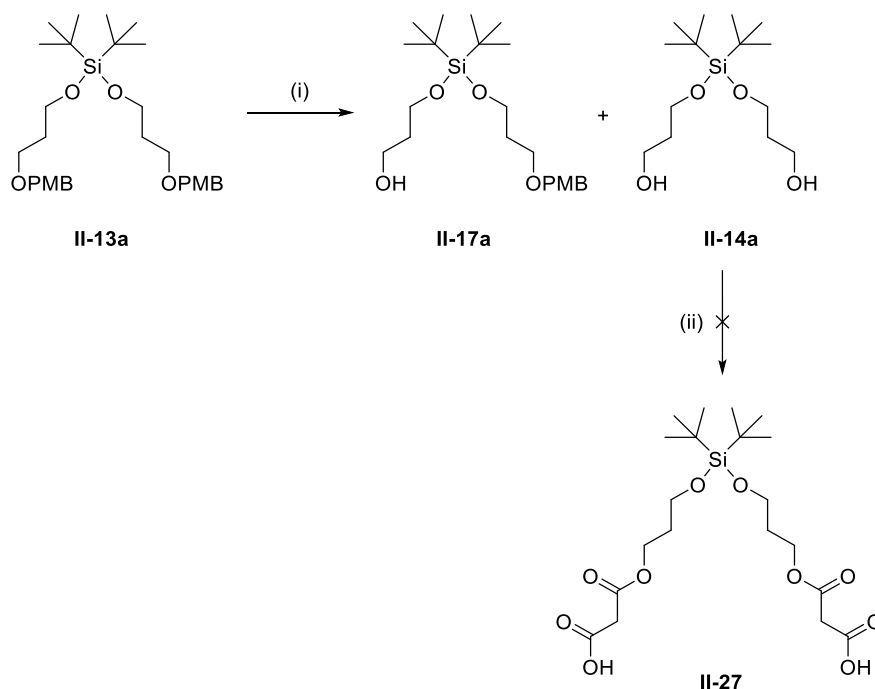


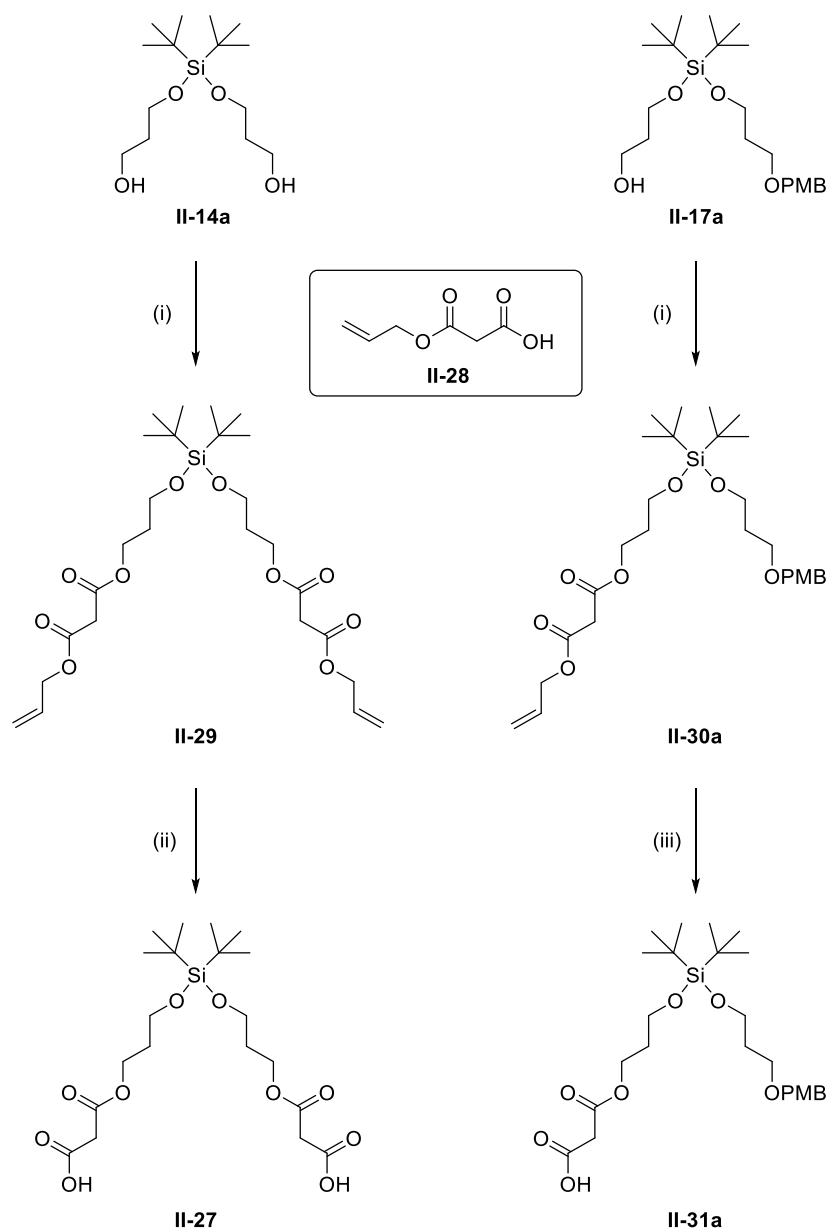
Figure II-11: Retrosynthetic analysis for the elaboration of cyclo-tris-malonate.

Treatment of **II-13** with 1 equiv. of DDQ provided the mono-protected derivative **II-17**, but also the diol **II-14** (Scheme II-8). The introduction of malonate moieties was initially attempted by reaction of **II-14** with Meldrum's acid at 120°C. However, the ether group was not stable under these conditions.



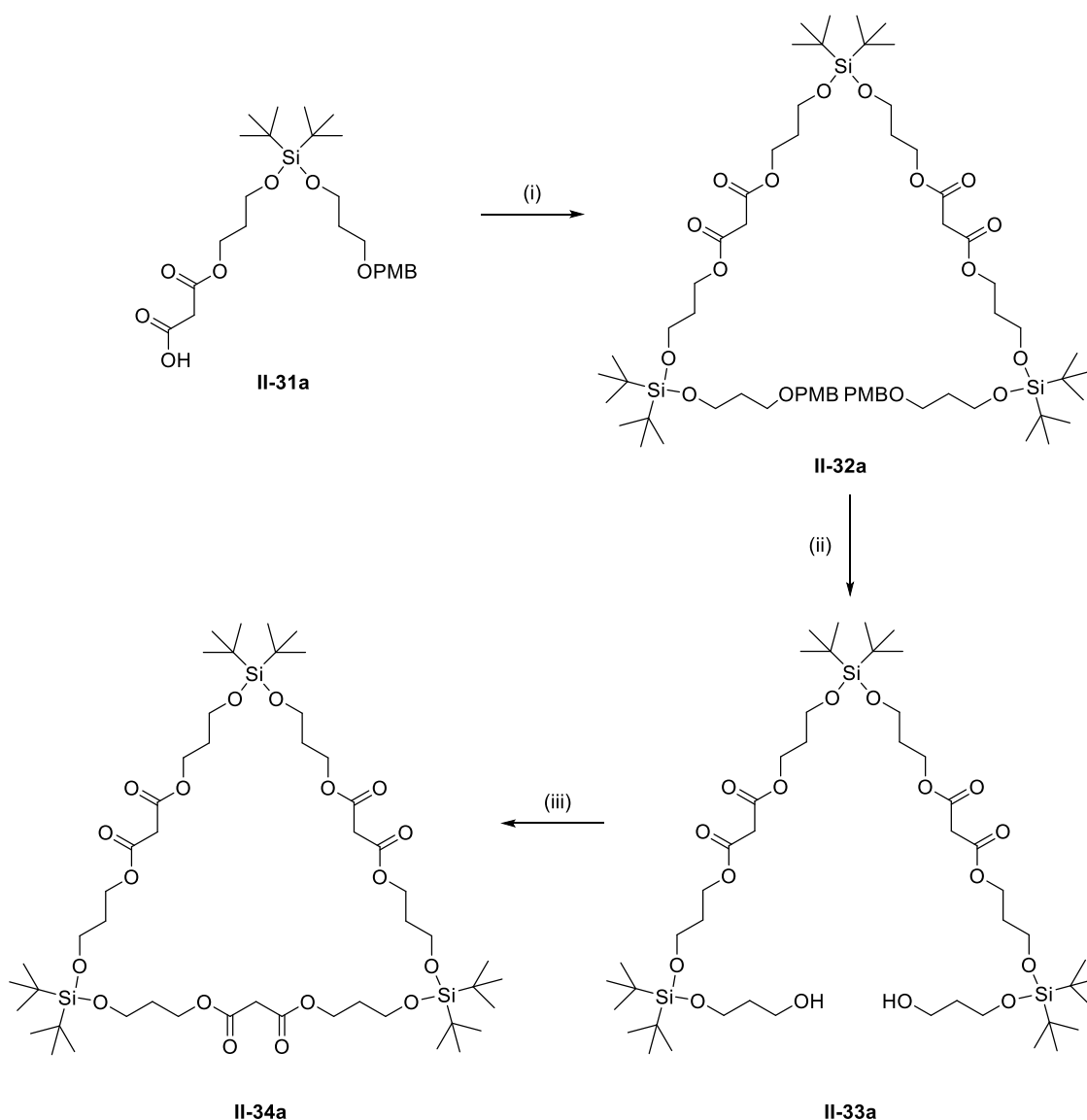
**Scheme II-8.** Reagents and conditions: (i) DDQ (1 eq), CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, rt, 1 h; (ii) Meldrum's acid, 120°C, (degradation).

In order to prevent the degradation of the starting material, an esterification reaction was carried out with **II-14** and malonic acid monoallyl ester **II-28** under the Steglich conditions (**Scheme II-9**).<sup>[15]</sup> The bis-protected derivative **II-29** was obtained in 74% yield. The esterification reaction was also performed with **II-17** to afford compound **II-30** in 73% yield. The choice of the appropriate protecting groups for the malonic acid functions was the key for this synthesis. The deprotection conditions must not be acidic or imply a fluorine reagent in order to preserve the bridging di-*t*-butylsilylene groups, may not be basic to preserve the ester functions and they should be inert toward the PMB protecting group. The allyl protecting groups were removed under neutral conditions by treatment with Pd(PPh<sub>3</sub>)<sub>4</sub> in the presence of an excess of morpholine.<sup>[16]</sup> Under these conditions the two other protecting groups (Si and PMB) and the ester functions remained effectively intact and the acidic derivatives **II-27** and **II-31** were obtained.



**Scheme II-9.** Reagents and conditions: (i) **II-28**, DCC, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, 0°C to rt, 12 h (**II-29**: 74%, **II-30a**: 73%); (ii) Pd(PPh<sub>3</sub>)<sub>4</sub>, piperidine, THF, rt, 4 h; (iii) Pd(PPh<sub>3</sub>)<sub>4</sub>, morpholine, THF, rt, 4 h.

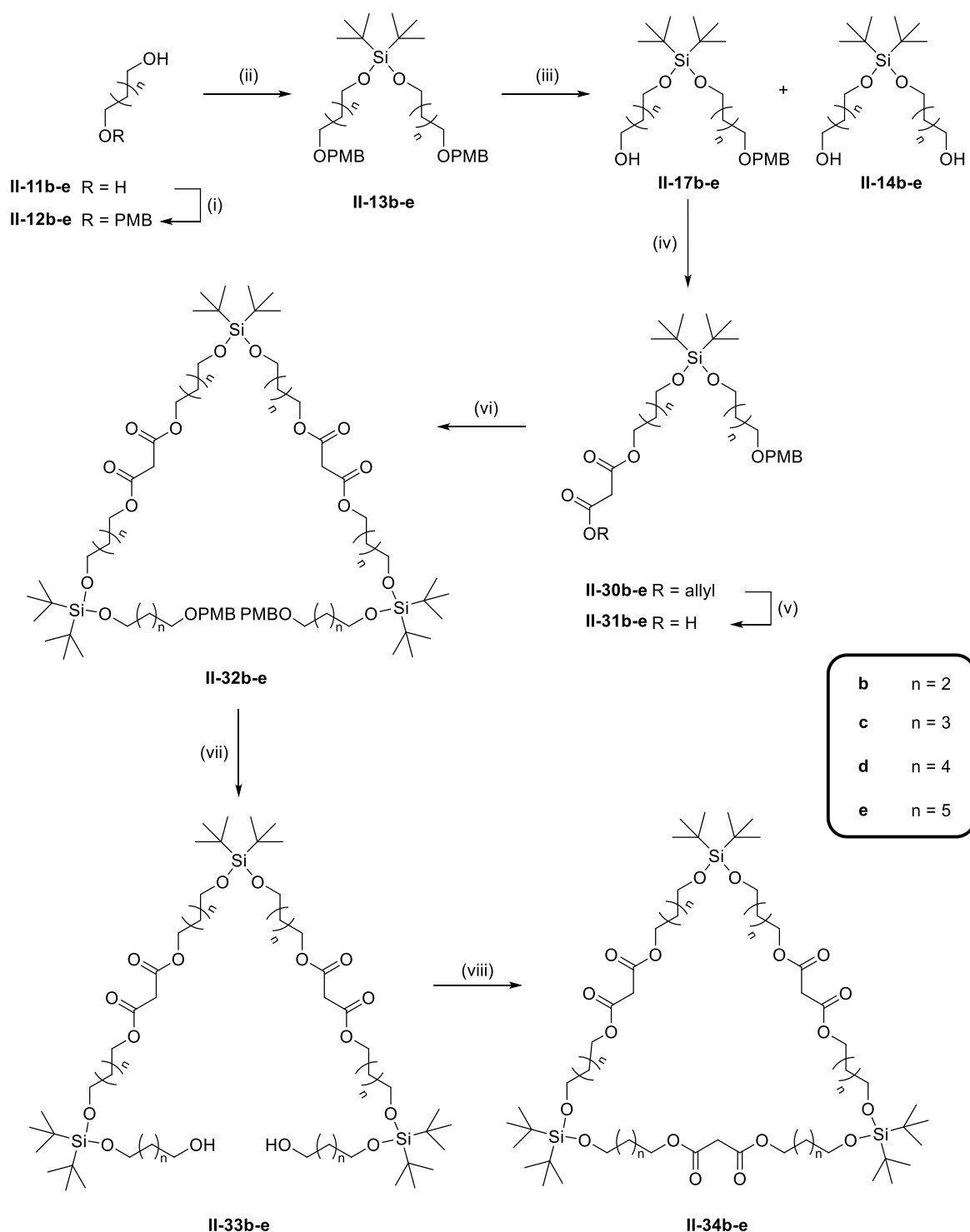
The bis-carboxylic acid derivative **II-27** being difficult to handle, only the mono-acidic derivative **II-31** was used (**Scheme II-10**). Treatment of **II-31a** with diol **II-14a** under esterification conditions using dicyclohexylcarbodiimide (DCC) and DMAP afforded **II-32a** in 61% yield. Subsequent deprotection of the PMB protecting group with DDQ gave the key product **II-33a** in a nearly quantitative yield (98%). Final macrocyclisation of **II-33a** with malonyl dichloride afforded the cyclo-tris-malonates **II-34a** in an excellent yield of 53%. It is also important to note that **II-34a** is the only formed cyclooligomer. By-product being polymers, the isolation of **II-34a** was easy to perform.



**Scheme II-10.** Reagents and conditions: (i) **II-14a**, DCC, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, 0°C to rt, 12 h (61%); (ii) DDQ, CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, rt, 6 h (98%); (iii) malonyl chloride, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, rt, 24 h (53%).

By following the same synthetic route, four others macrocyclic tris-malonates **II-34b-e** were synthesized starting from alkanediol possessing 4 to 7 carbon atoms (**Scheme II-11**). The 48-54-60 and 66 membered ring macrocycles **II-34b-e** were thus obtained in good yields (21-63%) considering their ring size.



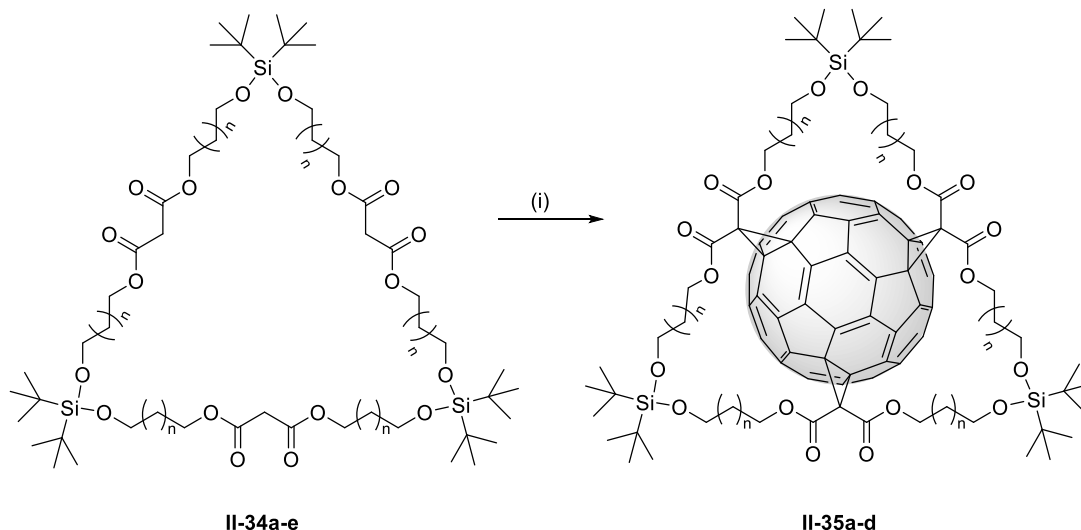


**Scheme II-11.** Reagents and conditions: (n= 2-5) (i) PMBCl, Ag<sub>2</sub>O, CH<sub>2</sub>Cl<sub>2</sub>, rt, 12 h; (ii) tBu<sub>2</sub>Si(OTf)<sub>2</sub>, DMF, pyridine, rt, 12 h (iii) DDQ (1 eq), CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, rt, 1 h; (iv) **II-28**, DCC, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, 0°C to rt, 12 h; (v) Pd(PPh<sub>3</sub>)<sub>4</sub>, morpholine, THF, rt, 4 h; (vi) **II-14b-e**, DCC, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, 0°C to rt, 12 h; (vii) DDQ, CH<sub>2</sub>Cl<sub>2</sub>, H<sub>2</sub>O, rt, 6 h; (viii) malonyl chloride, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, rt, 24 h.

3.2 Regioselective tris-functionalization of C<sub>60</sub>.

This work has been done in collaboration with Eric Meischner.

The reaction of the macrocyclic tris-malonates **II-34a-e** with C<sub>60</sub> was carried out in the presence of I<sub>2</sub> and DBU (**Scheme II-12**). The results are reported in [Table II-2](#). A poor regioselectivity was observed for the largest macrocycles. For the 66 membered macrocyclic tris-malonates **II-34e**, no C<sub>60</sub> tris-adducts could be isolated in a pure form from the mixture of products. In the case of **II-34c-d**, mixtures were also obtained but one specific regioisomer could be isolated in moderate yields. The large ring size and the relative flexibility of the O-Si-O motif bring a too high degree of freedom and consequently allows the formation of numerous tris-adducts and thus reduces the regioselectivity. For macrocycles **II-34a-b**, the reactions were highly regioselective and only the fullerene tris-adduct with an *e,e,e*-addition pattern was thus obtained. The relative position of the three cyclopropane rings on the C<sub>60</sub> were determined based on the C<sub>3</sub> symmetry deduced from the <sup>1</sup>H and <sup>13</sup>C NMR spectra. As shown in [Figure II-12](#), the 18 fullerene resonances of the sp<sup>2</sup> C atoms and the 2 fullerene resonances of the sp<sup>3</sup> C atoms observed in the <sup>13</sup>C NMR spectrum of **II-35a** are fully consistent with a threefold symmetrical compound. The cherry-red color and the UV-vis spectrum ([Figure II-9](#)) are also in complete agreement with the proposed *e,e,e*-addition pattern for compound **II-35**.



**Scheme II-12.** Reagents and conditions: (*n*= 1-5) (i) C<sub>60</sub>, I<sub>2</sub>, DBU, PhMe, rt, 1 h.

The *e* positions are the most reactive sites (kinetic products). Furthermore, the *e,e,e* tris-adducts of C<sub>60</sub> are the most stable regioisomers (thermodynamics products) for the macrocycles **II-34a-b** ([Table II-3](#)). The formation of the *e,e,e* tris-adducts of C<sub>60</sub> were optimal with the macrocycles **II-34a-b** and were thus obtained in excellent yields of 61% and 39%, respectively.

	(CH <sub>2</sub> ) <sub>3</sub> <b>II-34a</b>	(CH <sub>2</sub> ) <sub>4</sub> <b>II-34b</b>	(CH <sub>2</sub> ) <sub>5</sub> <b>II-34c</b>	(CH <sub>2</sub> ) <sub>6</sub> <b>II-34d</b>	(CH <sub>2</sub> ) <sub>7</sub> <b>II-34e</b>
e,e,e $C_3$	61	39	8	5	-
undefined $C_1$	-	-	35	26	-

Table II-2 : Results of the reactions of **II-34a-e** with  $C_{60}$  (%).

	e,e,e $C_3$	all-trans-4 $C_{3v}$	all-trans-3 $D_3$
(CH <sub>2</sub> ) <sub>3</sub> <b>II-34a</b>	0	+15	+115
(CH <sub>2</sub> ) <sub>4</sub> <b>II-34b</b>	0	+48	+26

Table II-3 : Calculated  $\Delta E$  heat of formation in kJ/mol at the AM1 semi-empirical level.

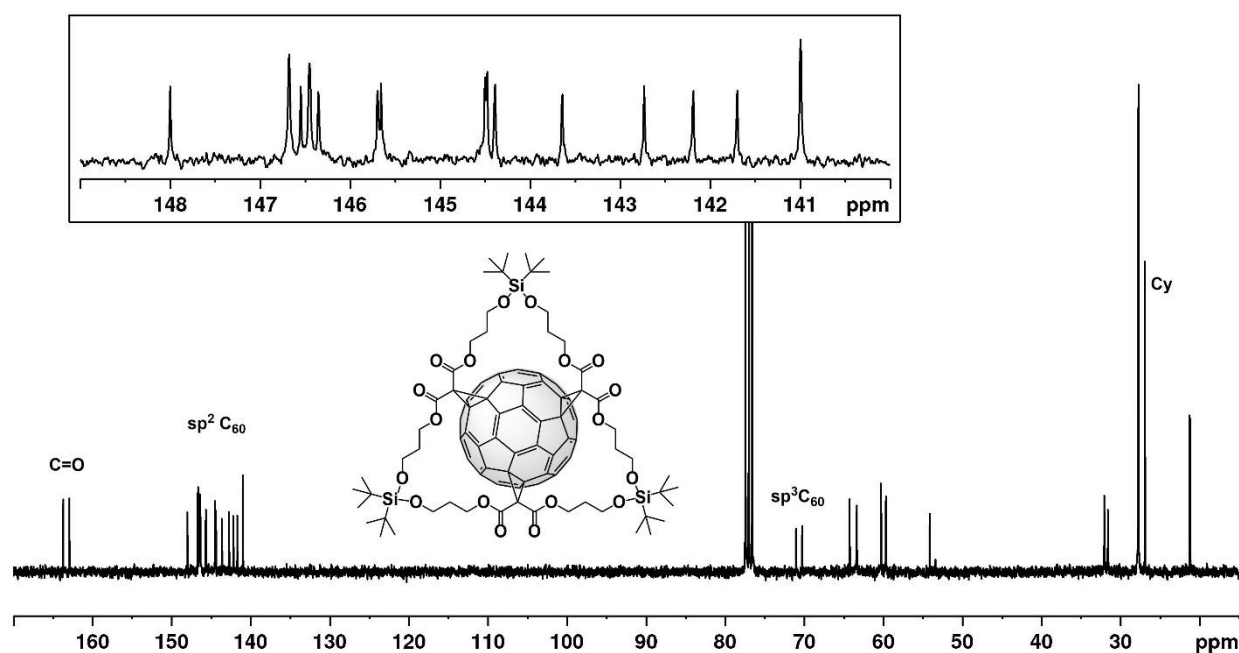


Figure II-12 : <sup>13</sup>C NMR spectrum (CDCl<sub>3</sub>, 100 MHz) of compound **II-34a** showing its  $C_3$  symmetry; Inset: detailed view showing the resonances of the fullerene sp<sup>2</sup> C atoms; Cy = cyclohexane.

## 4. Conclusion.

The reaction of macrocyclic di-*t*-butylsilylene-tethered bis- and tris-malonates with C<sub>60</sub> gave access to fullerene bis- and tris-adducts with an excellent regioselectivity. By the modification of the rigidity and the length of the spacer units linking the malonate subunits to the di-*t*-butylsilylene moieties, fullerene bis-adducts with different addition patterns were obtained.

The control of the regiochemistry is primarily governed by the ring size of the macrocycle. The O-Si-O motif play also an important role in the regioselectivity by introducing flexibility in the macrocycle. Beyond the flexibility/rigidity of the spacer, the regioselectivity is mainly influenced by the relative reactivity of the different double bonds of the mono-substituted fullerene intermediate. In most of the cases, the macrocyclisation on the fullerene core occurs under kinetic control. For this reason, mainly *e* and *trans-3* bis-adducts are formed in the case of the fullerene bis-adducts. Similarly, the fullerene tris-adducts with an all-*e* addition pattern are also preferentially formed owing to the higher reactivity of the equatorial positions.

The silyl groups are also protecting groups that can be readily cleaved to afford the corresponding acyclic fullerene polyols. This new strategy opens therefore new perspectives for the post-functionalization of fullerene multi-adducts derivatives. Furthermore, the obtained bis-adducts *e* and tris-adducts *e,e,e* are totally suited for the preparation of multi-functionalized C<sub>60</sub> hexa-adducts with an octahedral addition pattern.

## 5. Bibliography.

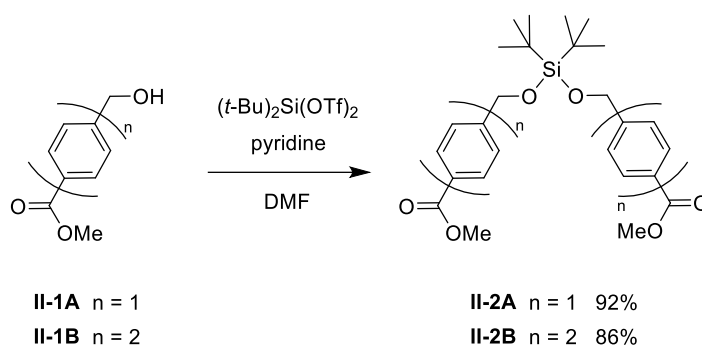
- [1] U. Reuther, T. Brandmüller, W. Donaubaue, F. Hampel, A. Hirsch, *Chem. – Eur. J.* **2002**, *8*, 2261–2273.
- [2] Z. Zhou, D. I. Schuster, S. R. Wilson, *J. Org. Chem.* **2003**, *68*, 7612–7617.
- [3] B. Gross, V. Schurig, I. Lamparth, A. Herzog, F. Djojo, A. Hirsch, *Chem. Commun.* **1997**, 1117–1118.
- [4] N. Chronakis, A. Hirsch, *Chem. Commun.* **2005**, 3709–3711.
- [5] M. Riala, N. Chronakis, *Org. Lett.* **2011**, *13*, 2844–2847.
- [6] D. Sigwalt, M. Holler, J.-F. Nierengarten, *Tetrahedron Lett.* **2013**, *54*, 3160–3163.
- [7] A. Bouzide, G. Sauv , *Tetrahedron Lett.* **1997**, *38*, 5945–5948.
- [8] J.-F. Nierengarten, T. Habicher, R. Kessinger, F. Cardullo, F. Diederich, V. Gramlich, J.-P. Gisselbrecht, C. Boudon, M. Gross, *Helv. Chim. Acta* **1997**, *80*, 2238–2276.
- [9] F. Djojo, A. Herzog, I. Lamparth, F. Hampel, A. Hirsch, *Chem. – Eur. J.* **1996**, *2*, 1537–1547.
- [10] W. Qian, Y. Rubin, *Angew. Chem. Int. Ed.* **1999**, *38*, 2356–2360.
- [11] T. Da Ros, M. Prato, V. Lucchini, *J. Org. Chem.* **2000**, *65*, 4289–4297.

- [12] Y. Nakamura, N. Takano, T. Nishimura, E. Yashima, M. Sato, T. Kudo, J. Nishimura, *Org. Lett.* **2001**, *3*, 1193–1196.
- [13] S. Sergeev, M. Schär, P. Seiler, O. Lukoyanova, L. Echegoyen, F. Diederich, *Chem. – Eur. J.* **2005**, *11*, 2284–2294.
- [14] A. Hirsch, I. Lamparth, H. R. Karfunkel, *Angew. Chem. Int. Ed. Engl.* **1994**, *33*, 437–438.
- [15] B. Neises, W. Steglich, *Angew. Chem.* **1978**, *90*, 556–557.
- [16] H. Kunz, H. Waldmann, *Angew. Chem. Int. Ed. Engl.* **1984**, *23*, 71–72.

## 6. Experimental part.

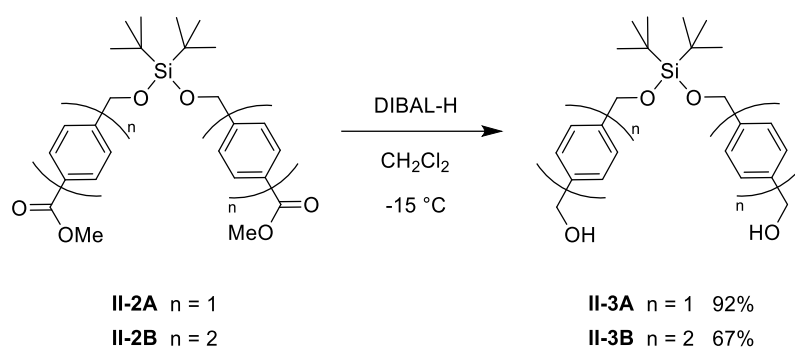
**General:** Reagents and solvents were purchased as reagent grade and used without further purification. All reactions were performed in standard glassware under an inert Ar atmosphere. Evaporation and concentration were done at water aspirator pressure and drying in vacuo at 10<sup>-2</sup> Torr. Column chromatography: silica gel 60 (230-400 mesh, 0.040-0.063 mm) was purchased from E. Merck. Thin Layer Chromatography (TLC) was performed on aluminum sheets coated with silica gel 60 F<sub>254</sub> purchased from E. Merck. IR spectra (cm<sup>-1</sup>) were recorded on a Perkin–Elmer Spectrum One Spectrophotometer. UV/Visible spectra were recorded on a Perkin-Elmer Lambda 35 UV/Vis Spectrometer. NMR spectra were recorded on a Bruker AC 300 or AC 400 with solvent peaks as reference at 25°C. MALDI-TOF-MS were obtained on a Bruker ULTRAFLEX TOF/TOF mass spectrometer with a dithranol matrix.

### Synthesis.



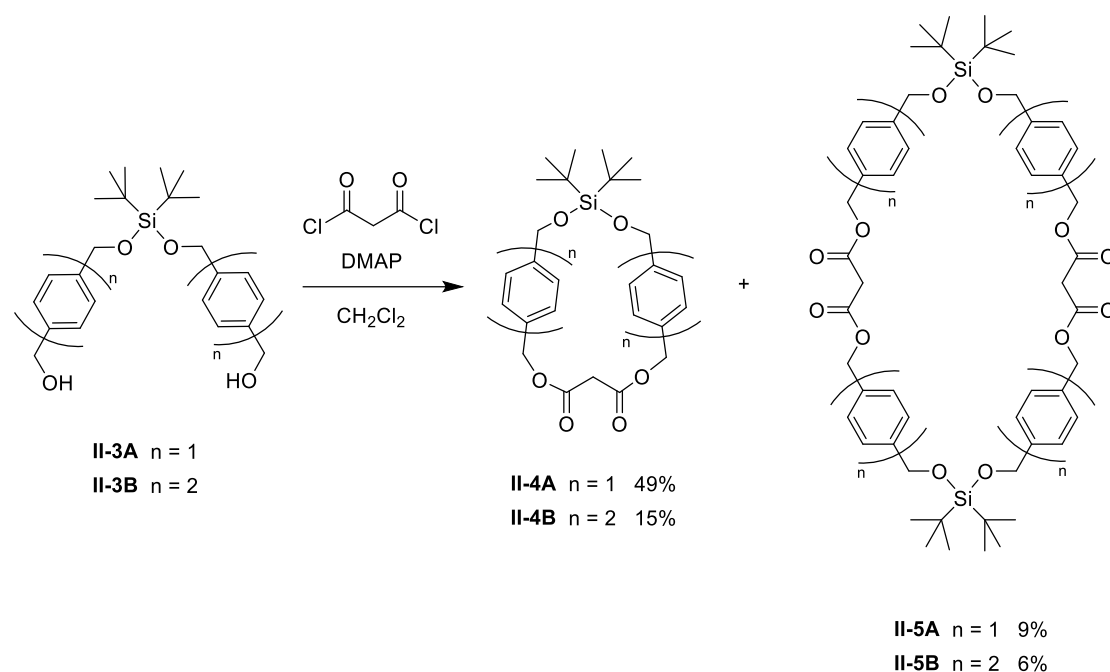
**Compound II-2A:** Di-tert-butylsilyl-bis(trifluoromethanesulfonate) (, 12.4mmol) was added slowly to a solution of **II-1A** (3.96 g, 24.8 mmol) and pyridine (2.1 mL, 26.1 mmol) in DMF (13 mL). The resulting mixture was stirred overnight at room temperature, then water was added and the aqueous layer extracted with Et<sub>2</sub>O. The organic layer was washed, dried (MgSO<sub>4</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2) yielded **II-2A** (4.94 g, 92%). IR (neat):  $\nu = 1723$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 8.00$  (d, J = 8 Hz, 4H), 7.36 (d, J = 8 Hz, 4H), 4.96 (s, 4H), 3.92 (s, 6H), 1.11 (s, 18H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 167.0, 146.2, 129.6, 128.9, 125.5, 65.2, 52.0, 27.9, 21.4$  ppm.

**Compound II-2B:** Di-tert-butylsilyl-bis(trifluoromethanesulfonate) (2.0 mL, 6.3 mmol) was added to a solution of **II-1B** (3.19 g, 13.2 mmol) and pyridine (1.1 mL, 13.5 mmol) in DMF (2 mL). The resulting mixture was stirred overnight at room temperature, then water was added and the aqueous layer extracted with CH<sub>2</sub>Cl<sub>2</sub>. The organic layer was washed, dried (MgSO<sub>4</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 7:3) yielded **II-2B** (3.31 g, 5.3 mmol, 86%) as a white solid. IR (neat):  $\nu = 1721$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 8.09$  (d,  $J = 8$  Hz, 4 H), 7.66 (d,  $J = 8$  Hz, 4 H), 7.60 (d,  $J = 8$  Hz, 4 H), 7.43 (d,  $J = 8$  Hz, 4 H), 5.01 (s, 4 H), 3.94 (s, 6 H), 1.14 (s, 18 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 167.0, 145.4, 141.3, 138.6, 130.1, 128.8, 127.1, 126.9, 126.3, 65.3, 52.1, 28.0, 21.4$  ppm.



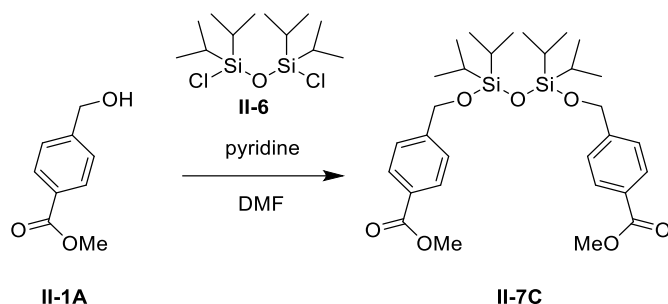
**Compound II-3A:** A solution of **II-2A** (2.42 g, 5.1 mmol) in dry CH<sub>2</sub>Cl<sub>2</sub> (20 mL) was added dropwise to a stirred 0.5 M DIBAL-H solution in dry CH<sub>2</sub>Cl<sub>2</sub> (22 mL, 22 mmol) at -15°C. The resulting mixture was stirred for 3 h, then MeOH and NH<sub>4</sub>Cl (aq) were carefully added. The resulting mixture was filtered (Celite) and evaporated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/Cyclohexane 9:1 to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 98:2) yielded **II-3A** (1.97 g, 92%) as a white solid. IR (neat):  $\nu = 3321$  (O-H) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.32$  (s, 8H), 4.93 (s, 4H), 4.68 (s, 4H), 1.10 (s, 18H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 140.7, 139.5, 127.0, 126.0, 65.4, 65.3, 28.0, 21.4$  ppm.

**Compound II-3B:** A solution of **II-2B** (3.03 g, 4.8 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (20 mL) was added dropwise to a stirred 1M DIBAL-H solution in THF (20.3 mL, 20.3 mmol) at -15°C. The resulting mixture was stirred for 3h, then quenched with MeOH and NH<sub>4</sub>Cl (aq), filtered on celite and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 96:4) yielded **II-3B** (1.85 g, 3.2 mmol, 67%) as a white solid. IR (neat):  $\nu = 3311$  (br, O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.59$  (d,  $J = 8$  Hz, 4 H), 7.56 (d,  $J = 8$  Hz, 4 H), 7.43 (d,  $J = 8$  Hz, 4 H), 7.41 (d,  $J = 8$  Hz, 4 H), 5.01 (s, 4 H), 4.74 (s, 4 H), 1.14 (s, 18 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 140.5, 140.4, 139.7, 139.4, 127.5, 127.2, 126.9, 126.3, 65.4, 65.1, 28.0, 21.4$  ppm.

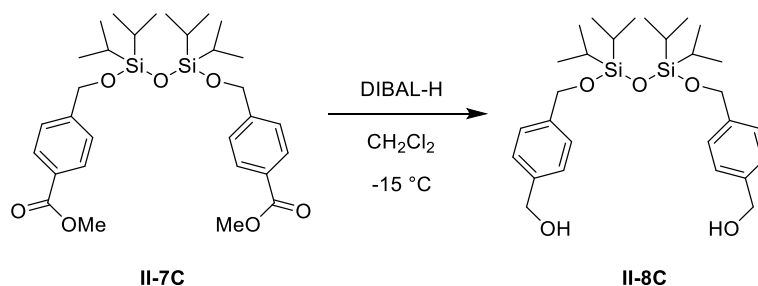


**Compound II-4A & II-5A:** A solution of malonyl chloride (0.5 mL, 5.2 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (100 mL) was added dropwise to a solution of **II-3A** (1.97 g, 4.7 mmol) and DMAP (1.27 g, 10.4 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (430 mL). The resulting mixture was stirred at room temperature for 2 h, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 9:1) yielded **II-4A** (1.12 g, 2.3 mmol, 49%) as a white solid and **II-5A** (0.20 g, 0.2 mmol, 9%) as a white solid. **II-4A:** IR (neat):  $\nu = 1732$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.01$  (s, 8 H), 5.13 (s, 4 H), 4.88 (s, 4 H), 3.51 (s, 2 H), 1.14 (s, 18 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.0, 141.0, 133.5, 127.3, 126.0, 66.8, 65.7, 42.4, 28.2, 21.3$  ppm. **II-5A:** <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.25$  (s, 16 H), 5.14 (s, 8 H), 4.87 (s, 8 H), 3.47 (s, 4 H), 1.10 (s, 36 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.2, 141.4, 133.8, 128.2, 125.9, 67.1, 65.2, 41.7, 28.0, 21.4$  ppm.

**Compound II-4B & II-5B:** A solution of malonyl chloride (0.32 mL, 3.2 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (100 mL) was added dropwise to a solution of **II-3B** (1.85 g, 3.2 mmol) and DMAP (1.47 g, 12 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (300 mL). The resulting mixture was stirred at room temperature for 2 h, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane/ether, 7:2:1) yielded **II-4B** (250 mg, 0.39 mmol, 12%) as a white solid and **II-5B** (170 mg, 0.13 mmol, 8%) as a white solid. **II-4B:** IR (neat):  $\nu = 1754$  (C=O), 1725 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.29$  (d,  $J = 8$  Hz, 4 H), 7.21 (d,  $J = 8$  Hz, 4 H), 7.19 (d,  $J = 8$  Hz, 4 H), 7.07 (d,  $J = 8$  Hz, 4 H), 5.22 (s, 4 H), 4.98 (s, 4 H), 3.56 (s, 2 H), 1.17 (s, 18 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.1, 140.7, 140.3, 138.5, 133.8, 128.2, 127.0, 126.4$  (two peaks), 67.0, 66.0, 42.3, 28.3, 21.5 ppm. **II-5B:** <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.54$ -7.46 (m, 16 H), 7.40-7.32 (m, 16 H), 5.21 (s, 8 H), 4.99 (s, 8 H), 3.53 (s, 4 H), 1.16 (s, 36 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.2, 141.1, 140.4, 139.1, 134.0, 128.9, 128.8, 127.1, 126.9, 126.3, 67.0, 65.4, 41.9, 28.1, 21.5$  ppm.

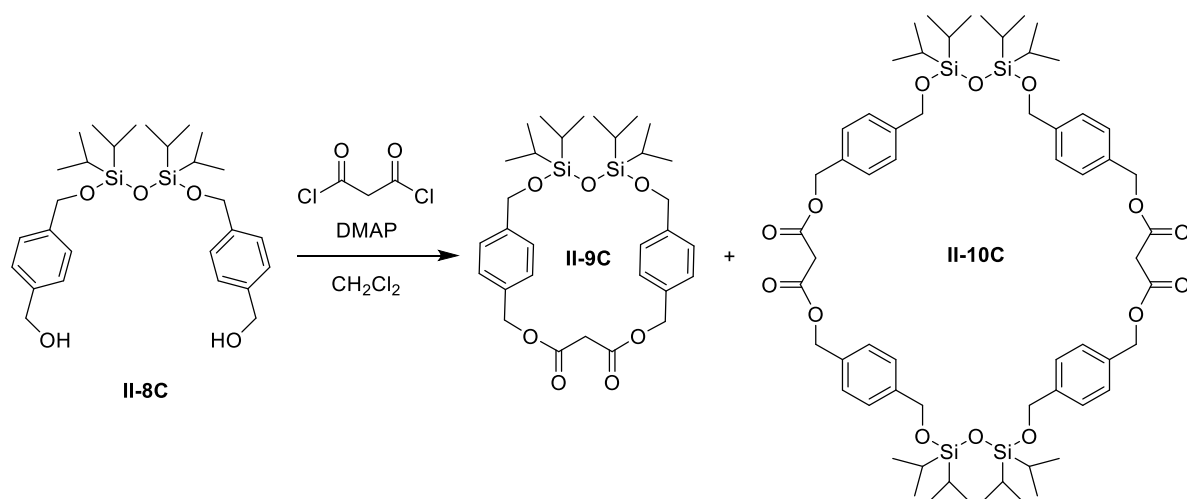


**Compound II-7C:** 1,3-Dichloro-1,1,3,3-tetraisopropylsiloxane (**II-6**) (2.47 mL, 7.9 mmol) was added to a solution of **II-1A** (2.77 g, 16.6 mmol) and pyridine (1.5 mL, 18.5 mmol) in DMF (9 mL). The mixture was stirred overnight at room temperature, then H<sub>2</sub>O was added and the product was extracted with ether and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2) yielded **II-7C** (4.53 g, 7.87 mmol, 99%) as a white solid. IR (neat):  $\nu = 1724$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.98$  (d,  $J = 8$  Hz, 4 H), 7.34 (d,  $J = 8$  Hz, 4 H), 4.88 (s, 4 H), 3.91 (s, 6 H), 1.06 (m, 28 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 167.0, 146.3, 129.5, 128.8, 125.5, 63.8, 52.0, 17.4, 17.3, 13.0$  ppm.

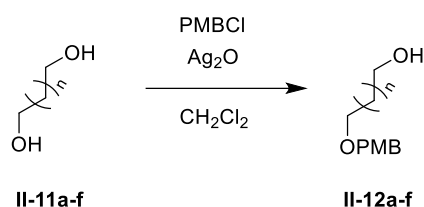


**Compound II-8C:** A solution of **II-7C** (4.51 g, 7.85 mmol) in dry THF (35 mL) was added dropwise to a stirred 0.5 M DIBAL-H solution in dry THF (33 mL, 33 mmol) at -15°C. The resulting mixture was stirred for 3 h, then MeOH and NH<sub>4</sub>Cl (aq) were carefully added. The resulting mixture was filtered (Celite) and evaporated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) yielded **II-8C** (3.16 g, 6.1 mmol, 77%) as a white solid. IR (neat):  $\nu = 3325$  (O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.27$  (s, 8 H), 4.84 (s, 4 H), 4.65 (s, 4 H), 1.08 (m, 28 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 140.7, 139.4, 126.9, 126.1, 65.2, 64.0, 17.4, 17.4, 13.1$  ppm.





**Compound II-9C & II-10C:** A solution of malonyl chloride (0.94 g, 6.6 mmol) in  $\text{CH}_2\text{Cl}_2$  (100 mL) was added dropwise to a solution of **II-8C** (3.14 g, 6.1 mmol) and DMAP (1.77g, 14.5 mmol) in  $\text{CH}_2\text{Cl}_2$  (590 mL). After 2 h, the resulting mixture was filtered on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$ ) and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 7:3 to  $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 9:1) yielded **II-9C** (2.59 g, 4.4 mmol, 73%) as a white solid and **II-10C** (170 mg, 0.14 mmol, 5%) as a white solid. **II-9C:** IR (neat):  $\nu = 1734$  (C=O)  $\text{cm}^{-1}$ .  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 7.13$  (d,  $J = 8$  Hz, 4 H), 7.04 (d,  $J = 8$  Hz, 4 H), 5.12 (s, 4 H), 4.85 (s, 4 H), 3.49 (s, 2 H), 1.11 (s, 28 H) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 75 MHz):  $\delta = 166.0$ , 141.1, 133.4, 127.8, 125.9, 67.0, 63.9, 42.3, 17.4 (two peaks), 13.1 ppm. **II-10C:**  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 7.22$  (m, 16 H), 5.12 (s, 8 H), 4.77 (s, 8 H), 3.44 (s, 4H), 1.07 (s, 56 H) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 166.3$ , 141.7, 133.7, 128.2, 125.9, 67.1, 63.8, 41.5, 17.4, 13.0 ppm.



a    n = 1    b    n = 2    c    n = 3    d    n = 4    e    n = 5    f    n = 6

**General Procedure for the mono-Protection of diols (II-12a-f) (GP I):** A solution of PMB-Cl (1eq.) in  $\text{CH}_2\text{Cl}_2$  was added dropwise to a solution of the appropriate diol (**II-11a-f**) (2-3 eq.) and  $\text{Ag}_2\text{O}$  (1 eq.) in  $\text{CH}_2\text{Cl}_2$ . The resulting mixture was stirred overnight at room temperature, then filtered on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 9:1) and concentrated. Column chromatography on  $\text{SiO}_2$  gave **II-12a-f**.

**Compound II-12a:** **II-12a** was synthesized from 1,3-propanediol (13.5 mL, 187 mmol), PMB-Cl (8.5 mL, 62.4 mmol) and  $\text{Ag}_2\text{O}$  (14.5 g, 62.4 mmol) in  $\text{CH}_2\text{Cl}_2$  (250 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 8:2 to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 9:1) yielded **II-12a** (9.77 g,

49.8 mmol, 80%) as a colorless oil. IR (neat):  $\nu = 3391$  (br, OH)  $\text{cm}^{-1}$ .  $^1\text{H-NMR}$  (400 MHz,  $\text{CDCl}_3$ ):  $\delta = 7.19$  (d,  $J = 8$  Hz, 2 H), 6.81 (d,  $J = 8$  Hz, 2 H), 4.39 (s, 2 H), 3.74 (s, 3 H), 3.71 (t,  $J = 6$  Hz, 2 H), 3.58 (t,  $J = 6$  Hz, 2 H), 1.78 (quint.,  $J = 6$  Hz, 2 H) ppm.  $^{13}\text{C-NMR}$  (100 MHz,  $\text{CDCl}_3$ ):  $\delta = 159.3$ , 130.1, 129.3, 113.9, 73.0, 69.3, 62.1, 55.3, 32.1 ppm.

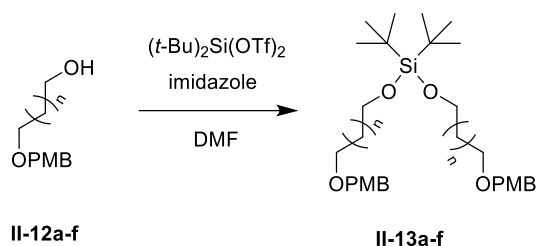
**Compound II-12b:** **II-12b** was synthesized from 1,4-butanediol (10.0 g, 111 mmol), PMB-Cl (5.0 mL, 37.0 mmol) and  $\text{Ag}_2\text{O}$  (8.6 g, 37.0 mmol) in  $\text{CH}_2\text{Cl}_2$  (150 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 8:2 to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 95:5) yielded **II-12b** (7.4 g, 35.1 mmol, 95%) as a colorless oil. IR (neat):  $\nu = 3392$  (br, OH)  $\text{cm}^{-1}$ .  $^1\text{H-NMR}$  (300 MHz,  $\text{CDCl}_3$ ):  $\delta = 7.25$  (d,  $J = 8$  Hz, 2 H), 6.87 (d,  $J = 8$  Hz, 2 H), 4.43 (s, 2 H), 3.80 (s, 3 H), 3.62 (t,  $J = 7$  Hz, 2 H), 3.45 (t,  $J = 7$  Hz, 4 H), 1.79 (m, 4 H) ppm.  $^{13}\text{C-NMR}$  (100 MHz,  $\text{CDCl}_3$ ):  $\delta = 159.2$ , 130.2, 129.3, 113.8, 72.7, 70.1, 62.7, 55.3, 30.2, 26.8 ppm.

**Compound II-12c:** **II-12c** was synthesized from 1,5-pentanediol (10.0 g, 96 mmol), PMB-Cl (4.4 mL, 32.0 mmol) and  $\text{Ag}_2\text{O}$  (7.4 g, 32.0 mmol) in  $\text{CH}_2\text{Cl}_2$  (150 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 8:2 to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 95:5) yielded **II-12c** (6.5 g, 29.1 mmol, 91%) as a colorless oil. IR (neat):  $\nu = 3392$  (br, OH)  $\text{cm}^{-1}$ .  $^1\text{H-NMR}$  (300 MHz,  $\text{CDCl}_3$ ):  $\delta = 7.25$  (d,  $J = 8$  Hz, 2 H), 6.87 (d,  $J = 8$  Hz, 2 H), 4.43 (s, 2 H), 3.80 (s, 3 H), 3.62 (t,  $J = 7$  Hz, 2 H), 3.45 (t,  $J = 7$  Hz, 4 H), 1.58 (m, 4 H), 1.43 (m, 2 H) ppm.  $^{13}\text{C-NMR}$  (100 MHz,  $\text{CDCl}_3$ ):  $\delta = 159.1$ , 130.7, 129.1, 113.7, 72.6, 70.0, 62.8, 55.3, 32.5, 29.5, 22.4 ppm.

**Compound II-12d:** **II-12d** was synthesized from 1,6-hexanediol (10.0 g, 85 mmol), PMB-Cl (3.8 mL, 28.2 mmol) and  $\text{Ag}_2\text{O}$  (6.6 g, 28.2 mmol) in  $\text{CH}_2\text{Cl}_2$  (200 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 93:7) yielded **II-12d** (6.2 g, 26.1 mmol, 92%) as a colorless oil. IR (neat):  $\nu = 3378$  (br, OH)  $\text{cm}^{-1}$ .  $^1\text{H-NMR}$  (400 MHz,  $\text{CDCl}_3$ ):  $\delta = 7.25$  (d,  $J = 8$  Hz, 2 H), 6.87 (d,  $J = 8$  Hz, 2 H), 4.43 (s, 2 H), 3.80 (s, 3 H), 3.62 (t,  $J = 7$  Hz, 2 H), 3.44 (t,  $J = 7$  Hz, 4 H), 1.58 (m, 4 H), 1.37 (m, 4 H) ppm.  $^{13}\text{C-NMR}$  (100 MHz,  $\text{CDCl}_3$ ):  $\delta = 159.1$ , 130.7, 129.2, 113.7, 72.6, 70.0, 63.0, 55.3, 32.7, 29.7, 26.0, 25.6 ppm.

**Compound II-12e:** **II-12e** was synthesized from 1,7-heptanediol (10.0 g, 76 mmol), PMB-Cl (4.1 mL, 30.3 mmol) and  $\text{Ag}_2\text{O}$  (7.7 g, 33.3 mmol) in  $\text{CH}_2\text{Cl}_2$  (200 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 95:5) yielded **II-12e** (6.9 g, 27.4 mmol, 90%) as a colorless oil. IR (neat):  $\nu = 3378$  (br, OH)  $\text{cm}^{-1}$ .  $^1\text{H-NMR}$  (400 MHz,  $\text{CDCl}_3$ ):  $\delta = 7.26$  (d,  $J = 8$  Hz, 2 H), 6.87 (d,  $J = 8$  Hz, 2 H), 4.43 (s, 2 H), 3.80 (s, 3 H), 3.63 (t,  $J = 7$  Hz, 2 H), 3.43 (t,  $J = 7$  Hz, 4 H), 1.58 (m, 4 H), 1.34 (m, 6 H) ppm.  $^{13}\text{C-NMR}$  (100 MHz,  $\text{CDCl}_3$ ):  $\delta = 159.1$ , 130.7, 129.2, 113.7, 72.5, 70.1, 63.0, 55.3, 32.7, 29.7, 29.2, 26.0, 25.7 ppm.

**Compound II-12f:** **II-12f** was synthesized from 1,8-octanediol (13.8 g, 95 mmol), PMB-Cl (4.3 mL, 31.6 mmol) and  $\text{Ag}_2\text{O}$  (8.0 g, 34.7 mmol) in  $\text{CH}_2\text{Cl}_2$  (270 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 95:5) yielded **II-12f** (5.8 g, 21.8 mmol, 69%) as a colorless oil. IR (neat):  $\nu = 3400$  (br, OH)  $\text{cm}^{-1}$ .  $^1\text{H-NMR}$  (400 MHz,  $\text{CDCl}_3$ ):  $\delta = 7.26$  (d,  $J = 8$  Hz, 2 H), 6.87 (d,  $J = 8$  Hz, 2 H), 4.43 (s, 2 H), 3.80 (s, 3 H), 3.63 (t,  $J = 7$  Hz, 2 H), 3.43 (t,  $J = 7$  Hz, 4 H), 1.57 (m, 4 H), 1.32 (m, 8 H) ppm.  $^{13}\text{C-NMR}$  (100 MHz,  $\text{CDCl}_3$ ):  $\delta = 159.1$ , 130.8, 129.2, 113.7, 72.5, 70.2, 63.1, 55.3, 32.8, 29.8, 29.4, 29.3, 26.1, 25.7 ppm.



**General Procedure for silylation of mono-protected diols (II-12a-f) (GP II):** Di-*tert*-butylsilyl-bis(trifluoromethanesulfonate) (1 eq.) was added slowly to a solution of **II-12a-f** (2 eq.) and pyridine (or imidazole) (2 eq.) in DMF. The resulting mixture was stirred overnight at room temperature, then water was added and the aqueous layer extracted with Et<sub>2</sub>O. The organic layer was washed, dried (MgSO<sub>4</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> gave **II-13a-f**.

**Compound II-13a:** **II-13a** was synthesized from **II-12a** (9.8 g, 49.8 mmol), (t-Bu)<sub>2</sub>Si(OTf)<sub>2</sub> (8.1 mL, 24.9 mmol) and imidazole (4.2 g, 62.2 mmol) in DMF (10 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded **II-13a** (9.0 g, 17.0 mmol, 68%) as a colorless oil. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.18 (d, *J* = 8 Hz, 4 H), 6.79 (d, *J* = 8 Hz, 4 H), 4.35 (s, 4 H), 3.84 (t, *J* = 6 Hz, 4 H), 3.72 (s, 6 H), 3.50 (t, *J* = 6 Hz, 4 H), 1.76 (quint., *J* = 6 Hz, 4 H), 0.91 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 159.1, 130.7, 129.6, 113.8, 72.7, 66.8, 60.7, 55.3, 33.1, 27.9, 21.2 ppm.

**Compound II-13b:** **II-13b** was synthesized from **II-12b** (7.3 g, 34.6 mmol), (t-Bu)<sub>2</sub>Si(OTf)<sub>2</sub> (5.1 mL, 15.7 mmol) and imidazole (2.4 g, 34.6 mmol) in DMF (20 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) yielded **II-13b** (5.3 g, 9.5 mmol, 60%) as a colorless oil. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.25 (d, *J* = 8 Hz, 4 H), 6.87 (d, *J* = 8 Hz, 4 H), 4.43 (s, 4 H), 3.81 (t, *J* = 6 Hz, 4 H), 3.80 (s, 6 H), 3.47 (t, *J* = 6 Hz, 4 H), 1.69 (m, 4 H), 1.61 (m, 4 H), 0.98 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 159.0, 130.7, 129.2, 113.7, 72.5, 70.0, 63.6, 55.3, 29.6, 27.9, 26.2, 21.1 ppm.

**Compound II-13c:** **II-13c** was synthesized from **II-12c** (5.6 g, 25.0 mmol), (t-Bu)<sub>2</sub>Si(OTf)<sub>2</sub> (3.7 mL, 11.4 mmol) and pyridine (2.0 mL, 25.0 mmol) in DMF (20 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) yielded **II-13c** (6.0 g, 10.3 mmol, 91%) as a colorless oil. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.25 (d, *J* = 8 Hz, 4 H), 6.87 (d, *J* = 8 Hz, 4 H), 4.43 (s, 4 H), 3.80 (s, 6 H), 3.79 (t, *J* = 7 Hz, 4 H), 3.45 (t, *J* = 7 Hz, 4 H), 1.59 (m, 4 H), 1.45 (m, 2 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 159.1, 130.8, 129.1, 113.7, 72.5, 70.2, 63.7, 55.3, 32.7, 29.6, 27.9, 22.5, 21.2 ppm.

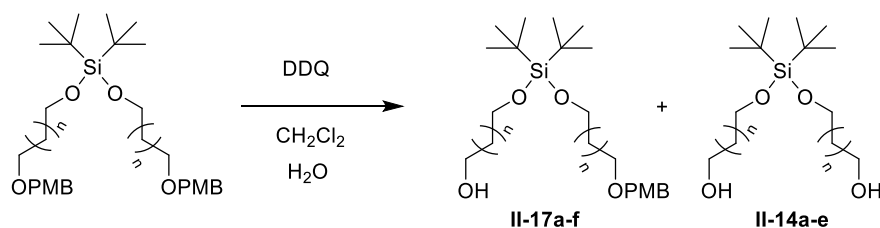
**Compound II-13d:** **II-13d** was synthesized from **II-12d** (6.4 g, 26.9 mmol), (t-Bu)<sub>2</sub>Si(OTf)<sub>2</sub> (4.0 mL, 12.3 mmol) and imidazole (2.2 g, 32.6 mmol) in DMF (20 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 10:3) yielded **II-13d** (4.0 g, 6.5 mmol, 59%) as a colorless oil. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.25 (d, *J* = 8 Hz, 4 H), 6.87 (d, *J* = 8 Hz, 4 H), 4.43 (s, 4 H), 3.80 (s, 6 H), 3.79 (t, *J* = 6 Hz, 4 H), 3.44 (t, *J* = 6 Hz, 4 H), 1.63 (m, 2 H), 1.58 (m, 4 H), 1.40 (m, 4 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 159.1, 130.8, 129.2, 113.7, 72.6, 70.2, 63.7, 55.3, 32.9, 29.9, 27.9, 26.1, 25.7, 21.1 ppm.

**Compound II-13e:** II-13e was synthesized from II-12e (6.3 g, 25.0 mmol), (t-Bu)<sub>2</sub>Si(OTf)<sub>2</sub> (5.0 mL, 11.4 mmol) and pyridine (1.9 mL, 23.8 mmol) in DMF (20 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) yielded II-13e (5.8 g, 9.0 mmol, 79%) as a colorless oil. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.27 (d, J = 8 Hz, 4 H), 6.87 (d, J = 8 Hz, 4 H), 4.43 (s, 4 H), 3.80 (s, 6 H), 3.79 (t, J = 6 Hz, 4 H), 3.43 (t, J = 6 Hz, 4 H), 1.56 (m, 8 H), 1.34 (m, 12 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 159.0, 130.8, 129.2, 113.7, 72.5, 70.2, 63.8, 55.3, 32.9, 29.8, 29.3, 27.9, 26.3, 25.8, 21.2 ppm.

**Compound II-13f:** II-13f was synthesized from II-12f (5.8 g, 21.8 mmol), (t-Bu)<sub>2</sub>Si(OTf)<sub>2</sub> (3.2 mL, 9.9 mmol) and pyridine (1.7 mL, 20.8 mmol) in DMF (20 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) yielded II-13f (5.5 g, 8.2 mmol, 83%) as a colorless oil. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.25 (d, J = 8 Hz, 4 H), 6.87 (d, J = 8 Hz, 4 H), 4.43 (s, 4 H), 3.80 (s, 6 H), 3.79 (t, J = 6 Hz, 4 H), 3.43 (t, J = 6 Hz, 4 H), 1.55 (m, 8 H), 1.32 (m, 16 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 159.0, 130.8, 129.2, 113.7, 72.5, 70.3, 63.8, 55.3, 32.9, 29.9, 29.5, 29.4, 27.9, 26.2, 25.8, 21.2 ppm.

**Compound II-14a:** A solution of II-13a (2.54 g, 4.8 mmol) and DDQ (2.27 g, 10.0 mmol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (170/4 mL) was stirred at room temperature for 6 h, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 96:4) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 98:2) yielded II-14a (1.19 g, 4.1 mmol, 86%) as a colorless oil. IR (neat): ν = 3326 (br, OH) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 3.98 (t, J = 6 Hz, 4 H), 3.76 (t, J = 6 Hz, 4 H), 1.98 (br s, 2 H), 1.78 (quint., J = 6 Hz, 4 H), 0.97 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 62.5, 61.1, 34.7, 27.9, 21.1 ppm.

**Compound II-14d:** A solution of II-13d (4.00 g, 6.5 mmol) and DDQ (3.68 g, 16.2 mmol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (180/10 mL) was stirred overnight at room temperature, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 9:1) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 94:6) yielded II-14d (2.00 g, 5.3 mmol, 82%) as a colorless oil. IR (neat): ν = 3325 (br, OH) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 3.81 (t, J = 6 Hz, 4 H), 3.65 (t, J = 6 Hz, 4 H), 1.58 (m, 8 H), 1.40 (m, 8 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 63.7, 63.0, 32.9, 32.8, 27.9, 25.6, 25.5, 21.2 ppm.



**General procedure for the synthesis of compounds II-14a-e & II-17a-f:** Water (20 eq) was added to a solution of II-13a-f (1 eq) and 2,3-dichloro-5,6-dicyano-1,4-benzoquinone (DDQ, 1 eq) in CH<sub>2</sub>Cl<sub>2</sub> at room temperature. After 1 h, the crude was filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) and concentrated. Column chromatography on SiO<sub>2</sub> gave II-14a-e and II-17a-f.

**Compound II-14a & II-17a:** II-14a and II-17a were synthesized from water (6 mL), II-13a (6.56 g, 12.3 mmol) and DDQ (2.80 g, 12.3 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (450 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) gave II-17a (2.22 g, 44%) and II-14a (0.82 g, 2.8 mmol, 23%) as colorless oils. II-17a: IR (neat):  $\nu = 3448$  (O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.25$  (d,  $J = 8$  Hz, 2 H), 6.87 (d,  $J = 8$  Hz, 2 H), 4.43 (s, 2 H), 4.00 (t,  $J = 6$  Hz, 2 H), 3.95 (t,  $J = 6$  Hz, 2 H), 3.80 (s, 3 H), 3.79 (t,  $J = 6$  Hz, 2 H), 1.86 (m,  $J = 6$  Hz, 2 H), 1.78 (m,  $J = 6$  Hz, 2 H), 1.62 (s br, 1 H), 1.01 (s, 18 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 159.2, 130.6, 129.3, 113.8, 72.7, 66.5, 62.6, 61.4, 60.9, 55.3, 34.6, 33.0, 27.8, 21.1$  ppm.

**Compound II-14b & II-17b:** II-14b and II-17b were synthesized from water (3 mL), II-13b (6.00 g, 10.7 mmol) and DDQ (2.43 g, 10.7 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (270 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 9:1 to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) gave II-17b (2.22 g, 5.0 mmol, 47%) and II-14b (0.75 g, 2.3 mmol, 22%) as colorless oils. II-14b: IR (neat):  $\nu = 3327$  (O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 3.88$  (t,  $J = 6$  Hz, 4 H), 3.68 (t,  $J = 6$  Hz, 4 H), 2.00 (s br, 2 H), 1.68 (m, 8 H), 1.01 (s, 18 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 64.0, 62.9, 29.9, 29.7, 28.0, 21.3$  ppm. II-17b: IR (neat):  $\nu = 3400$  (O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.26$  (t,  $J = 8.5$  Hz, 2 H), 6.87 (t,  $J = 8.5$  Hz, 2 H), 4.44 (s, 2 H), 3.85 (m, 4 H), 3.80 (s, 3 H), 3.66 (t,  $J = 6$  Hz, 2 H), 3.48 (t,  $J = 6$  Hz, 2 H), 1.66 (m, 9 H), 1.00 (s, 18 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 159.3, 130.9, 129.4, 113.9, 72.7, 70.1, 63.9, 63.9, 63.0, 55.4, 30.0, 29.8, 29.7, 28.0, 26.4, 21.3$  ppm.

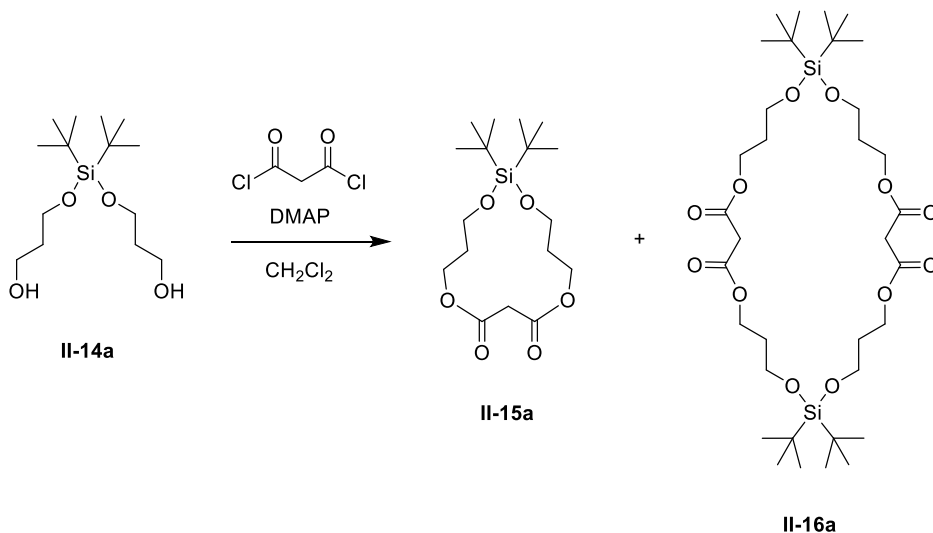
**Compound II-14c & II-17c:** II-14c and II-17c were synthesized from water (3 mL), II-13c (6.32 g, 10.7 mmol) and DDQ (2.44 g, 10.7 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (270 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 9:1 to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) gave II-17c (2.38 g, 5.1 mmol, 47%) and II-14c (0.77 g, 2.2 mmol, 21%) as colorless oils. II-14c: IR (neat):  $\nu = 3327$  (O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 3.83$  (t,  $J = 6$  Hz, 4 H), 3.65 (t,  $J = 6$  Hz, 4 H), 1.58 (m, 8 H), 1.45 (m, 4 H), 1.00 (s, 18 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 63.8, 63.1, 32.8, 32.6, 28.0, 22.2, 21.3$  ppm. II-17c: IR (neat):  $\nu = 3399$  (O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.25$  (d,  $J = 8$  Hz, 2 H), 6.87 (d,  $J = 8$  Hz, 2 H), 4.43 (s, 2 H), 3.81 (m, 7 H), 3.63 (t,  $J = 7$  Hz, 2 H), 3.44 (t,  $J = 7$  Hz, 2 H), 1.59 (m, 8 H), 1.43 (m, 4 H), 1.00 (s, 18 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 159.1, 130.7, 129.2, 113.7, 72.5, 70.2, 63.7, 63.6, 62.9, 55.3, 32.8, 32.6, 32.5, 29.6, 27.9, 22.5, 22.0, 21.2$  ppm.

**Compound II-14d & II-17d:** II-14d and II-17d were synthesized from water (3 mL), II-13d (4.30 g, 7.0 mmol) and DDQ (1.58 g, 7.0 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (250 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) gave II-17d (1.51 g, 3.0 mmol, 43%) and II-14d (0.58 g, 1.5 mmol, 22%) as colorless oils. II-14d: IR (neat):  $\nu = 3325$  (br, OH) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 3.81$  (t,  $J = 6$  Hz, 4 H), 3.65 (t,  $J = 6$  Hz, 4 H), 1.58 (m, 8 H), 1.40 (m, 8 H) 0.99 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 63.7, 63.0, 32.9, 32.8, 27.9, 25.6, 25.5, 21.2$  ppm. II-17d: IR (neat):  $\nu = 3399$  (O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.26$  (d,  $J = 8.5$  Hz, 2 H), 6.87 (d,  $J = 8.5$  Hz, 2 H), 4.43 (s, 2 H), 3.80 (m, 7 H), 3.63 (t,  $J = 6.5$  Hz, 2 H), 3.44 (t,  $J = 6.5$  Hz, 2 H), 1.57 (m, 8 H), 1.38 (m, 9 H), 0.99 (s,

18 H) ppm. <sup>13</sup>C NMR (75 MHz, CDCl<sub>3</sub>): δ = 159.2, 130.9, 129.4, 113.9, 72.7, 70.3, 63.9, 63.8, 63.0, 55.4, 33.0, 33.0, 32.9, 29.9, 28.0, 26.2, 25.9, 25.8, 25.7, 21.3 ppm.

**Compound II-14e & II-17e:** **II-14e** and **II-17e** were synthesized from water (3 mL), **II-13e** (5.00 g, 7.8 mmol) and DDQ (1.76 g, 7.8 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (200 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 9:1 to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 97:3) gave **II-17e** (2.08 g, 3.9 mmol 51%) and **II-14e** (0.65 g, 1.6 mmol, 21%) as colorless oils. **II-14e**: IR (neat): ν = 3327 (O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): δ = 3.80 (t, *J* = 6.5 Hz, 4H), 3.64 (t, *J* = 6.5 Hz, 4 H), 1.56 (m, 8 H), 1.39 (m, 12 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>): δ = 63.9, 63.2, 33.0, 32.9, 29.4, 28.0, 26.0, 25.9, 21.3 ppm. **II-17e**: IR (neat): ν = 3387 (O-H) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.26 (d, *J* = 8.5 Hz, 2 H), 6.88 (d, *J* = 8.5 Hz, 2 H), 4.43 (s, 2 H), 3.80 (m, 7 H), 3.63 (t, *J* = 6.5 Hz, 2 H), 3.43 (t, *J* = 6.5 Hz, 2 H), 1.55 (m, 8 H), 1.36 (m, 12 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>): δ = 159.3, 130.9, 129.4, 113.9, 72.7, 70.4, 64.0, 63.9, 63.2, 55.4, 33.0, 33.0, 32.9, 29.9, 29.5, 29.4, 28.0, 26.4, 26.0, 25.9, 21.3 ppm.

**Compound II-17f:** **II-17f** was synthesized from water (15 mL), **II-13f** (5.4 g, 8.1 mmol) and DDQ (1.8 g, 8.1 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (270 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 95:5) yielded **II-17f** (2.3 g, 4.1 mmol, 50%) as a colorless oil. IR (neat): ν = 3388 (br, OH) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.27 (d, *J* = 8 Hz, 2 H), 6.87 (d, *J* = 8 Hz, 2 H), 4.43 (s, 2 H), 3.80 (m, 7 H), 3.63 (t, *J* = 6 Hz, 2 H), 3.43 (t, *J* = 6 Hz, 2 H), 1.55 (m, 8 H), 1.33 (m, 16 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 159.1, 130.7, 129.2, 113.7, 72.5, 70.2, 63.8, 63.1, 55.3, 32.9, 32.8, 29.8, 29.5, 29.4 (two peaks), 27.9, 27.5, 26.2, 25.8, 25.7, 21.2 ppm.



**Compounds II-15a & II-16a:** A solution of DMAP (1.42 g, 11.6 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (100 mL) was added dropwise to a solution of **II-14a** (1.56 g, 5.3 mmol) and malonyl chloride (0.57 mL, 5.8 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (480 mL). The resulting mixture was stirred overnight at room temperature, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded **II-15a** (0.45 g, 1.3 mmol, 24%) as a colorless oil, **II-16a** (0.29 g, 0.4 mmol, 15%) as a colorless oil. **II-15a**: IR (neat): ν = 1735 (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 4.29 (t, *J* = 6 Hz, 4 H), 3.86 (t, *J* = 6 Hz, 4 H), 3.37 (s, 2 H), 1.91 (quint., *J* =



3.47 (t,  $J = 6$  Hz, 4 H), 3,37 (s, 2 H), 1.77 – 1.59 (m, 16 H), 0.97 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.1, 159.1, 130.7, 129.2, 113.7, 72.5, 70.0, 65.5, 63.6, 63.1, 55.3, 41.6, 29.6, 29.1, 27.9, 26.3, 25.15, 21.2$  ppm.

**Compound II-18c:** II-18c was synthesized from II-17c (2.3 g, 4.8 mmol), malonyl chloride (0.23 mL, 2.4 mmol) and DMAP (0.65 g, 5.3 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (100 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 9:1) yielded II-18c (2.3 g, 2.3 mmol, 94%) as a colorless oil. IR (neat):  $\nu = 1753$  (C=O),  $1736$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.25$  (d,  $J = 8$  Hz, 4 H), 6.87 (d,  $J = 8$  Hz, 4 H), 4.43 (s, 4 H), 4.14 (t,  $J = 7$  Hz, 4 H), 3.80 (s, 6 H), 3.79 (t,  $J = 7$  Hz, 8 H), 3.45 (t,  $J = 7$  Hz, 4 H), 3.35 (s, 2 H), 1.69 – 1.50 (m, 16 H), 1.38 (m, 8 H), 0.99 (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.6, 159.0, 130.7, 129.1, 113.7, 72.5, 70.2, 65.6, 63.7, 63.4, 55.3, 41.6, 32.9, 32.7, 29.8, 28.5, 27.9, 26.1, 25.7, 25.6, 25.5, 21.2$  ppm.

**Compound II-18d:** II-18d was synthesized from II-17d (1.2 g, 2.3 mmol), malonyl chloride (0.16 g, 1.2 mmol) and DMAP (0.31 g, 2.6 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (120 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 95:5) yielded II-18d (1.1 g, 1.1 mmol, 93%) as a colorless oil. IR (neat):  $\nu = 1753$  (C=O),  $1736$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.25$  (d,  $J = 8$  Hz, 4 H), 6.87 (d,  $J = 8$  Hz, 4 H), 4.43 (s, 4 H), 4.14 (t,  $J = 7$  Hz, 4 H), 3.80 (s, 6 H), 3.79 (t,  $J = 6$  Hz, 8 H), 3.44 (t,  $J = 6$  Hz, 4 H), 3.36 (s, 2 H), 1.69 – 1.50 (m, 22 H), 1.38 (m, 16 H), 0.99 (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.6, 159.0, 130.7, 129.2, 113.7, 72.5, 70.2, 65.6, 63.8, 63.6, 55.3, 41.6, 32.7, 32.4, 29.6, 28.3, 27.9, 22.5, 22.1, 21.2$  ppm.

**Compound II-18e:** II-18e was synthesized from II-17e (2.3 g, 4.3 mmol), malonyl chloride (0.31 g, 2.2 mmol) and DMAP (0.58 g, 4.7 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (150 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 98:2) yielded II-18e (2.3 g, 2.1 mmol, 96%) as a colorless oil. IR (neat):  $\nu = 1753$  (C=O),  $1736$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.26$  (d,  $J = 8$  Hz, 4 H), 6.87 (d,  $J = 8$  Hz, 4 H), 4.43 (s, 4 H), 4.13 (t,  $J = 7$  Hz, 4 H), 3.80 (s, 6 H), 3.79 (t,  $J = 6$  Hz, 8 H), 3.43 (t,  $J = 6$  Hz, 4 H), 3.36 (s, 2 H), 1.66 – 1.51 (m, 16 H), 1.35 (m, 24 H), 0.99 (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.6, 159.1, 130.8, 129.1, 113.7, 72.5, 70.2, 65.6, 63.8, 63.7, 55.3, 41.6, 32.9, 32.8, 29.8, 29.3, 29.0, 28.5, 28.0, 26.3, 25.8, 25.7, 21.2$  ppm.

**Compound II-18f:** II-18f was synthesized from II-17f (2.3 g, 4.1 mmol), malonyl chloride (0.29 g, 2.0 mmol) and DMAP (1.1 g, 9.0 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (150 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 98:2) yielded II-18f (2.1 g, 1.8 mmol, 90%) as a colorless oil. IR (neat):  $\nu = 1753$  (C=O),  $1736$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.25$  (d,  $J = 8$  Hz, 4 H), 6.87 (d,  $J = 8$  Hz, 4 H), 4.43 (s, 4 H), 4.13 (t,  $J = 7$  Hz, 4 H), 3.80 (s, 6 H), 3.79 (t,  $J = 6$  Hz, 8 H), 3.43 (t,  $J = 6$  Hz, 4 H), 3.36 (s, 2 H), 1.66 – 1.50 (m, 16 H), 1.32 (m, 32 H), 0.99 (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.7, 159.0, 130.8, 129.2, 113.7, 72.5, 70.2, 65.7, 63.8$  (two peaks), 55.3, 41.7, 32.9, 29.8, 29.5, 29.4, 29.3, 29.2, 28.7, 27.5, 26.2, 25.8, 21.2 ppm.







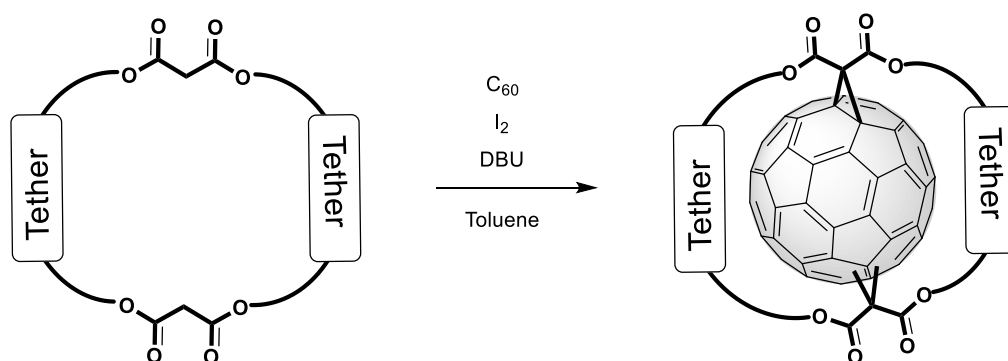
777.38 (24%, [M+H]<sup>+</sup>, calcd. for C<sub>38</sub>H<sub>73</sub>O<sub>12</sub>Si<sub>2</sub>: 777.46), 815.38 (20%, [M+K]<sup>+</sup>, calcd. for C<sub>38</sub>H<sub>72</sub>O<sub>12</sub>Si<sub>2</sub>K: 815.42).

**Compound II-20c:** **II-20c** was synthesized from **II-19c** (1.40 g, 1.83 mmol), malonyl chloride (0.2 mL, 2.01 mmol) and DMAP (0.27 g, 2.20 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (160 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 97:3) yielded **II-20c** (430 mg, 0.52 mmol, 28%) as a colorless oil. IR (neat):  $\nu = 1752$  (C=O),  $1734$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.15$  (t,  $J = 7$  Hz, 8 H),  $3.81$  (t,  $J = 7$  Hz, 8 H),  $3.36$  (s, 4 H),  $1.68$  (quint,  $J = 7$  Hz, 8 H),  $1.58$  (quint.,  $J = 7$  Hz, 8 H),  $1.45$  (m, 16 H),  $0.99$  (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.6$ ,  $65.5$ ,  $63.5$ ,  $41.7$ ,  $32.4$ ,  $28.3$ ,  $27.9$ ,  $22.1$ ,  $21.2$  ppm. MALDI-TOF-MS:  $833.37$  (100%, [M+H]<sup>+</sup>, calcd. for C<sub>42</sub>H<sub>81</sub>O<sub>12</sub>Si<sub>2</sub>: 833.53),  $855.51$  (61%, [M+Na]<sup>+</sup>, calcd. for C<sub>42</sub>H<sub>80</sub>O<sub>12</sub>Si<sub>2</sub>Na: 855.51).

**Compound II-20d:** **II-20d** was synthesized from **II-19d** (660 mg, 0.80 mmol), malonyl chloride (0.09 mL, 0.88 mmol) and DMAP (118 mg, 0.96 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (80 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 98:2) yielded **II-20d** (212 mg, 0.24 mmol, 30%) as a colorless oil. IR (neat):  $\nu = 1753$  (C=O),  $1735$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.15$  (t,  $J = 7$  Hz, 8 H),  $3.80$  (t,  $J = 7$  Hz, 8 H),  $3.36$  (s, 4H),  $1.66$  (quint,  $J = 7$  Hz, 8 H),  $1.56$  (quint.,  $J = 7$  Hz, 8 H),  $1.39$  (m, 16 H),  $0.99$  (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.6$ ,  $65.5$ ,  $63.6$ ,  $41.8$ ,  $32.7$ ,  $28.5$ ,  $27.9$ ,  $25.6$ ,  $25.4$ ,  $21.2$  ppm. MALDI-TOF-MS:  $889.73$  ([M+H]<sup>+</sup>, calcd. for C<sub>46</sub>H<sub>89</sub>O<sub>12</sub>Si<sub>2</sub>: 889.59).

**Compound II-20e:** **II-20e** was synthesized from **II-19e** (1.47 g, 1.68 mmol), malonyl chloride (0.17 mL, 1.76 mmol) and DMAP (0.43 g, 3.52 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (150 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 98:2) yielded **II-20e** (298 mg, 0.32 mmol, 19%) as a colorless oil. IR (neat):  $\nu = 1752$  (C=O),  $1736$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.14$  (t,  $J = 7$  Hz, 8 H),  $3.80$  (t,  $J = 7$  Hz, 8 H),  $3.36$  (s, 4 H),  $1.65$  (quint,  $J = 7$  Hz, 8 H),  $1.55$  (m, 8 H),  $1.35$  (m, 24 H),  $1.00$  (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.6$ ,  $65.6$ ,  $63.7$ ,  $41.8$ ,  $32.8$ ,  $29.0$ ,  $28.4$ ,  $25.8$ ,  $25.7$ ,  $21.1$  ppm. MALDI-TOF-MS:  $967.68$  (100%, [M+Na]<sup>+</sup>, calcd. for C<sub>50</sub>H<sub>96</sub>O<sub>12</sub>Si<sub>2</sub>Na: 967.63),  $983.60$  (28%, [M+K]<sup>+</sup>, calcd. for C<sub>50</sub>H<sub>96</sub>O<sub>12</sub>Si<sub>2</sub>K: 983.61),  $945.57$  (26%, [M]<sup>+</sup>, calcd. for C<sub>50</sub>H<sub>96</sub>O<sub>12</sub>Si<sub>2</sub>: 945.65).

**Compound II-20f:** **II-20f** was synthesized from **II-19f** (1.30 g, 1.39 mmol), malonyl chloride (0.15 mL, 1.53 mmol) and DMAP (0.37 g, 3.06 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (200 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 98:2) yielded **II-20f** (287 mg, 0.29 mmol, 19%) as a colorless oil. IR (neat):  $\nu = 1752$  (C=O),  $1736$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.14$  (t,  $J = 7$  Hz, 8 H),  $3.80$  (t,  $J = 7$  Hz, 8 H),  $3.36$  (s, 4 H),  $1.73$  (quint,  $J = 7$  Hz, 8 H),  $1.64$  (quint.,  $J = 6$  Hz, 8 H),  $1.55$  (m, 32 H),  $1.00$  (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.6$ ,  $65.6$ ,  $63.8$ ,  $41.8$ ,  $32.9$ ,  $29.3$ ,  $29.2$ ,  $28.5$ ,  $27.9$ ,  $25.7$ ,  $25.7$ ,  $21.2$  ppm. MALDI-TOF-MS:  $1023.72$  (100%, [M+Na]<sup>+</sup>, calcd. for C<sub>54</sub>H<sub>104</sub>O<sub>12</sub>Si<sub>2</sub>Na: 1023.70),  $1039.69$  (30%, [M+K]<sup>+</sup>, calcd. for C<sub>54</sub>H<sub>104</sub>O<sub>12</sub>Si<sub>2</sub>K: 1039.67),  $1000.63$  (12%, [M]<sup>+</sup>, calcd. for C<sub>54</sub>H<sub>104</sub>O<sub>12</sub>Si<sub>2</sub>: 1000.71).



**General Procedure for the formation of bis-adducts :** DBU (5 eq.) was added to a solution of C<sub>60</sub> (1 eq.), macrocycle bis-malonate (1 eq.) and I<sub>2</sub> (2.5 eq.) in toluene (2 mL/mg of C<sub>60</sub>). The resulting mixture was stirred at room temperature for 1 h, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> gave **II-21** and/or **II-22** and/or **II-23**.

**Compound II-21A:** **II-21A** was synthesized from **II-5A** (190 mg, 0.2 mmol), C<sub>60</sub> (141 mg, 0.2 mmol), I<sub>2</sub> (124 mg, 0.5 mmol) and DBU (0.15 mL, 1.0 mmol) in toluene (280 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5) yielded **II-21A** (179 mg, 0.1 mmol, 54%) as a brown glassy solid. IR (neat):  $\nu = 1748$  (C=O),  $1729$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 252$  (6x10<sup>4</sup>), 322 (sh, 2x10<sup>4</sup>), 413 (2x10<sup>3</sup>), 423 (2x10<sup>3</sup>), 493 (1x10<sup>3</sup>), 579 (sh, 6x10<sup>2</sup>), 635 (sh, 3x10<sup>2</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.47$  (d, <sup>3</sup>J = 8 Hz, 4 H), 7.26 (d, <sup>3</sup>J = 8 Hz, 8 H), 7.13 (d, <sup>3</sup>J = 8 Hz, 4 H), 5.78 (d, <sup>2</sup>J = 11 Hz, 2 H), 5.46 (d, <sup>2</sup>J = 11 Hz, 2 H), 5.35 (d, <sup>2</sup>J = 11 Hz, 2 H), 5.33 (d, <sup>2</sup>J = 11 Hz, 2 H), 4.91 (d, <sup>2</sup>J = 14 Hz, 2 H), 4.88 (d, <sup>2</sup>J = 14 Hz, 2 H), 4.77 (d, <sup>2</sup>J = 14 Hz, 2 H), 4.72 (d, <sup>2</sup>J = 14 Hz, 2 H), 1.07 (s, 18 H), 1.06 (s, 18 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.8, 163.2, 146.7, 146.5, 146.3$  (two peaks), 145.7, 145.3, 145.2, 145.1, 144.8, 144.6, 144.5, 144.4, 144.1, 143.8, 143.6, 143.2, 143.0, 142.9, 142.6, 142.3, 142.1, 142.0, 141.8, 141.6, 141.3, 141.0, 139.3, 138.6, 137.9, 133.3, 132.6, 130.6, 130.3, 125.6 (two peaks), 70.9 (two peaks), 69.4, 68.8, 65.3, 65.1, 51.3, 27.9 (two peaks), 26.9, 21.6, 21.4 ppm. MS-MALDI-TOF: 1686.7 ([M]<sup>+</sup>, calcd for C<sub>114</sub>H<sub>69</sub>O<sub>12</sub>Si<sub>2</sub>: 1686.4).

**Compound II-21B & II-22B & II-23B:** **II-21B & II-22B & II-23B** were synthesized from **II-5B** (170 mg, 0.14 mmol), C<sub>60</sub> (100 mg, 0.14 mmol), I<sub>2</sub> (89 mg, 0.35 mmol) and DBU (0.1 mL, 0.7 mmol) in toluene (200 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5) yielded **II-21B** (40 mg, 20  $\mu$ mol, 15%) as a brown glassy solid, **II-22B** (12 mg, 6  $\mu$ mol, 4%) as a brown glassy solid and **II-23B** (16 mg, 8  $\mu$ mol, 6%) as a brown glassy solid. **II-21B:** IR (neat):  $\nu = 1749$  (C=O),  $1728$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max} = 253, 317$  (sh), 412, 423, 489 nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.57$  (d, <sup>3</sup>J = 8 Hz, 4 H), 7.52 (d, <sup>3</sup>J = 8 Hz, 4 H), 7.41 (s, 8 H), 7.38 (d, <sup>3</sup>J = 8 Hz, 4 H), 7.33 (d, <sup>3</sup>J = 8 Hz, 4 H), 7.21 (d, <sup>3</sup>J = 8 Hz, 4 H), 7.13 (d, <sup>3</sup>J = 8 Hz, 4 H), 5.73 (d, <sup>2</sup>J = 12 Hz, 2 H), 5.63 (d, <sup>2</sup>J = 12 Hz, 2 H), 5.49 (d, <sup>2</sup>J = 12 Hz, 2 H), 5.40 (d, <sup>2</sup>J = 12 Hz, 2 H), 4.83 (d, <sup>2</sup>J = 14 Hz, 2 H), 4.80 (d, <sup>2</sup>J = 14 Hz, 2 H), 4.71 (d, <sup>2</sup>J = 13 Hz, 2 H), 4.65 (d, <sup>2</sup>J = 13 Hz, 2 H), 1.14 (s, 18 H), 1.10 (s, 18 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.5, 163.4, 146.8, 146.6, 146.2, 146.1, 145.1, 145.8, 145.7, 145.6, 145.5, 145.4, 145.1, 145.0, 144.4, 144.1, 143.7, 143.5, 143.3, 143.2,$

143.1, 142.5, 142.2, 142.0, 141.9, 141.8 (two peaks), 141.6, 141.3, 140.5, 140.0, 139.3, 139.0, 138.8, 137.8, 133.5, 133.2, 130.4, 129.9, 127.4, 127.1, 126.9, 126.8 (two peaks), 126.4, 71.4, 70.9, 69.1, 68.9, 65.5, 65.3, 51.7, 28.2, 28.1, 21.3, 21.2 ppm. MS-MALDI-TOF: 1990.3 ([M+H]<sup>+</sup>, calcd for C<sub>138</sub>H<sub>85</sub>O<sub>12</sub>Si<sub>2</sub>: 1990.5). **II-22B**: IR (neat):  $\nu = 1747$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max} = 256, 311$  (sh), 398 (sh), 409 (sh), 422, 479 nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.53$  (d, <sup>3</sup>J = 8 Hz, 2 H), 7.49 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.48-7.45 (m, 4 H), 7.44 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.41 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.33 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.30 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.29 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.24 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.18 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.16 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.04 (d, <sup>3</sup>J = 8 Hz, 2 H), 6.99 (d, <sup>3</sup>J = 8 Hz, 2 H), 5.85 (d, <sup>2</sup>J = 11 Hz, 1 H), 5.77 (d, <sup>2</sup>J = 12 Hz, 1 H), 5.60 (d, <sup>2</sup>J = 12 Hz, 1 H), 5.59 (d, <sup>2</sup>J = 11 Hz, 1 H), 5.47 (d, <sup>2</sup>J = 12 Hz, 1 H), 5.33 (d, <sup>2</sup>J = 11 Hz, 1 H), 5.17 (d, <sup>2</sup>J = 11 Hz, 1 H), 5.07 (d, <sup>2</sup>J = 12 Hz, 1 H), 5.03 (s, 2 H), 4.84 (d, <sup>2</sup>J = 13 Hz, 1 H), 4.74 (d, <sup>2</sup>J = 13 Hz, 1 H), 4.68 (s, 2 H), 4.45 (d, <sup>2</sup>J = 13 Hz, 1 H), 4.34 (d, <sup>2</sup>J = 13 Hz, 1 H), 1.15 (s, 18 H), 1.10 (s, 18 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.9, 163.7, 163.4, 162.6, 147.6, 147.4, 147.1$  (two peaks), 146.4, 146.3 (two peaks), 146.2, 146.0, 145.9, 145.8, 145.5 (two peaks), 145.4, 145.2, 145.1, 145.0, 144.6 (two peaks), 144.5 (three peaks), 144.4, 144.3 (three peaks), 144.0 (two peaks), 143.7 (two peaks), 143.6, 143.4, 143.3, 143.2 (two peaks), 143.0 (two peaks), 142.9, 142.8, 142.7, 142.5, 142.3, 142.2, 142.0, 141.9, 141.7, 141.6, 141.5, 141.4, 141.3, 141.0, 140.8, 140.7, 140.6, 140.5 (two peaks), 139.4, 138.8, 138.7, 138.4, 138.3, 138.1, 133.7, 133.4 (two peaks), 133.1, 130.9, 130.2, 129.0, 128.9, 127.5, 127.4, 127.3, 127.2, 127.1, 126.9, 126.8, 126.6 (two peaks), 126.4, 125.9, 71.4 (three peaks), 69.0, 68.9, 68.7, 68.6, 66.2, 65.7, 65.3, 65.1, 51.0, 48.3, 28.3, 28.2, 28.1 (two peaks), 21.4, 21.2 (two peaks) ppm. **II-23B**: IR (neat):  $\nu = 1746$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max} = 412$  (sh), 438, 471 nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.42$  (d, <sup>3</sup>J = 8 Hz, 4 H), 7.28 (d, <sup>3</sup>J = 8 Hz, 4 H), 7.20 (s, 8 H), 7.18 (d, <sup>3</sup>J = 8 Hz, 4 H), 7.09 (d, <sup>3</sup>J = 8 Hz, 4 H), 7.06 (d, <sup>3</sup>J = 8 Hz, 4 H), 6.95 (d, <sup>3</sup>J = 8 Hz, 4 H), 5.70 (d, <sup>2</sup>J = 12 Hz, 2 H), 5.69 (d, <sup>2</sup>J = 12 Hz, 2 H), 5.35 (d, <sup>2</sup>J = 12 Hz, 2 H), 5.03 (s, 4 H), 4.90 (d, <sup>2</sup>J = 12 Hz, 2 H), 4.83 (d, <sup>2</sup>J = 13 Hz, 2 H), 4.74 (d, <sup>2</sup>J = 13 Hz, 2 H), 1.23 (s, 9 H), 1.21 (s, 9 H), 1.18 (s, 9 H), 1.15 (s, 9 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.3, 162.4, 147.4, 146.2, 145.9$  (two peaks), 145.7, 145.4, 145.3 (two peaks), 145.1, 145.0, 144.7, 144.6, 144.4, 144.2, 143.8, 143.6, 143.2, 142.5, 142.0, 141.0 (two peaks), 140.9, 140.5, 140.2, 139.4, 138.7 (two peaks), 138.4, 138.2, 137.5, 133.5, 133.4, 129.5, 128.3, 127.1, 127.0, 126.6 (three peaks), 126.5, 70.1, 68.6, 68.0, 67.7, 66.0, 65.6, 49.0, 28.5, 28.3, 28.2 (two peaks), 21.6, 21.5, 21.4, 21.2 ppm.

**Compound II-21C & II-22C:** **II-21C & II-22C** were synthesized from **II-10C** (170 mg, 0.14 mmol), C<sub>60</sub> (104 mg, 0.14 mmol), I<sub>2</sub> (89 mg, 0.35 mmol) and DBU (0.1 mL, 0.7 mmol) in toluene (210 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5) yielded **II-21C** (42 mg, 22  $\mu$ mol, 15%) as a brown glassy solid and **II-22C** (72 mg, 38  $\mu$ mol, 26%) as a brown glassy solid. **II-21C**: IR (neat):  $\nu = 1751$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 254$  (1.3x10<sup>5</sup>), 311 (sh, 5x10<sup>4</sup>), 397, 409 (sh), 422, 481 nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.41$  (d, <sup>3</sup>J = 8 Hz, 2 H), 7.35 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.33 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.28 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.26 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.21 (d, <sup>3</sup>J = 8 Hz, 2 H), 7.04 (d, <sup>3</sup>J = 8 Hz, 2 H), 6.98 (d, <sup>3</sup>J = 8 Hz, 2 H), 5.73 (d, <sup>2</sup>J = 11 Hz, 1 H), 5.67 (d, <sup>2</sup>J = 11

Hz, 1 H), 5.61 (d, <sup>2</sup>J = 12 Hz, 1 H), 5.49 (d, <sup>2</sup>J = 12 Hz, 1 H), 5.37 (d, <sup>2</sup>J = 12 Hz, 1 H), 5.19 (d, <sup>2</sup>J = 12 Hz, 1 H), 5.11 (d, <sup>2</sup>J = 11 Hz, 1 H), 5.00 (d, <sup>2</sup>J = 11 Hz, 1 H), 4.84 (s, 2 H), 4.77 (s, 2 H), 4.65 (d, <sup>2</sup>J = 13 Hz, 1 H), 4.58 (d, <sup>2</sup>J = 13 Hz, 1 H), 4.56 (s, 2 H), 1.06-0.80 (m, 56 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz): δ = 163.6, 163.5, 163.3, 162.6, 148.1, 147.3, 147.2, 147.1, 146.4 (two peaks), 146.2, 146.1, 145.9, 145.5 (two peaks), 145.3, 145.2, 144.9, 144.8, 144.7, 144.6 (two peaks), 144.5 (two peaks), 144.4, 144.3 (three peaks), 144.0, 143.9, 143.8, 143.7, 143.6, 143.5, 143.4, 143.3 (two peaks), 143.1 (two peaks), 142.9, 142.8 (two peaks), 142.5, 142.3, 142.2, 142.1, 142.0 (three peaks), 141.9, 141.8, 141.5 (two peaks), 141.3, 141.0, 140.7, 138.9, 138.8, 138.3, 138.2, 133.9, 133.3, 132.7, 132.5, 129.9, 129.0, 128.7, 128.5, 126.7, 126.1, 125.9, 125.5, 71.4, 71.3 (two peaks), 69.7, 68.9, 68.8 (two peaks), 68.7, 64.4, 63.9, 63.8, 63.5, 53.7, 51.0, 17.4 (two peaks), 17.3 (four peaks), 17.2 (three peaks), 13.1, 13.0 (two peaks), 12.9 (two peaks), 12.8 ppm. MS-MALDI-TOF: 1889.3 ([M]<sup>+</sup>, calcd for C<sub>122</sub>H<sub>88</sub>O<sub>14</sub>Si<sub>4</sub>: 1889.5). **II-22C**: IR (neat): ν = 1747 (C=O), 1728 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>): λ<sub>max</sub>(ε) = 251 (1.3x10<sup>5</sup>), 319 (sh, 5x10<sup>4</sup>), 399 (sh), 412, 423, 473, 489 nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): δ = 7.44 (d, <sup>3</sup>J = 8 Hz, 4 H, ArH), 7.28 (d, <sup>3</sup>J = 8 Hz, 4 H, ArH), 7.26 (d, <sup>3</sup>J = 8 Hz, 4 H, ArH), 7.14 (d, <sup>3</sup>J = 8 Hz, 4 H), 5.77 (d, <sup>2</sup>J = 11 Hz, 2 H), 5.43 (d, <sup>2</sup>J = 11 Hz, 2 H), 5.40 (d, <sup>2</sup>J = 11 Hz, 2 H), 5.30 (d, <sup>2</sup>J = 11 Hz, 2 H), 4.80 (d, <sup>2</sup>J = 13 Hz, 2 H), 4.76 (d, <sup>2</sup>J = 13 Hz, 2 H), 4.72 (d, <sup>2</sup>J = 13 Hz, 2 H), 4.68 (d, <sup>2</sup>J = 13 Hz, 2 H), 1.05-0.97 (m, 56 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz): δ = 163.8, 163.3, 146.8, 146.3 (three peaks), 145.8, 145.4, 145.3, 145.2, 145.1, 145.0, 144.9, 144.6 (two peaks), 144.4, 144.1, 144.0, 143.8, 143.6, 143.4, 143.2, 143.1, 143.0, 142.8, 142.6, 142.5 (two peaks), 142.1, 142.0, 141.9, 141.7, 141.4, 141.2, 140.7, 139.6, 138.8 (two peaks), 137.9, 133.3, 132.9, 132.7, 130.9, 130.3, 129.8, 126.1, 126.0, 125.9, 71.0, 70.9, 69.3, 68.9, 64.1, 63.9, 51.4, 17.4 (two peaks), 17.3, 13.0, 12.9 (two peaks) ppm. MS-MALDI-TOF: 1890.3 ([M+H]<sup>+</sup>, calcd for C<sub>122</sub>H<sub>89</sub>O<sub>14</sub>Si<sub>4</sub>: 1890.5).

**Compounds II-23a:** **II-23a** was synthesized from **II-16a** (220 mg, 0.3 mmol), C<sub>60</sub> (219 mg, 0.3 mmol), I<sub>2</sub> (190 mg, 0.8 mmol) and DBU (0.21 mL, 1.4 mmol) in toluene (440 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5 to CH<sub>2</sub>Cl<sub>2</sub>) yielded **II-23a** (115 mg, 0.08 mmol, 27%) as a brown glassy solid. IR (neat): ν = 1748 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>): λ<sub>max</sub>(ε = 259 (84200), 323 (sh, 24700), 439 (2500), 476 (2100), 622 (sh, 250) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 4.71-4.62 (m, 4 H), 4.50-4.41 (m, 4 H), 4.03 (m, 4 H), 3.88 (m, 4 H), 2.10 (m, 4 H), 1.98 (m, 4 H), 1.11 (s, 9 H), 1.04 (s, 18 H), 0.87 (s, 9 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 163.5, 163.3, 148.2, 147.5, 147.4 (two peaks), 147.1, 146.3, 146.0, 145.8, 145.7, 145.5, 145.3, 145.05, 145.0, 144.8, 144.5 (two peaks), 144.4, 144.2, 143.7 (two peaks), 143.6, 142.5, 142.45, 141.4, 141.1, 140.3, 136.4, 136.0, 135.9, 135.7, 70.6, 67.69, 64.0, 63.5, 60.4, 59.9, 50.2, 32.3, 31.9, 28.0, 27.95, 27.85, 27.8, 21.3, 21.1, 21.05 ppm. MALDI-TOF-MS: 1438.4 ([M+H]<sup>+</sup>, calcd for C<sub>94</sub>H<sub>61</sub>O<sub>12</sub>Si<sub>2</sub>: 1438.4).

**Compound II-23b:** **II-23b** was synthesized from **II-20b** (286 mg, 0.37 mmol), C<sub>60</sub> (265 mg, 0.37 mmol), I<sub>2</sub> (234 mg, 0.92 mmol) and DBU (0.25 mL, 1.7 mmol) in toluene (530 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:2 to 4:1) followed by gel permeation

chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) yielded **II-23b** (212 mg, 0.14 mmol, 39%) as a brown glassy solid. IR (neat):  $\nu = 1746$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 259$  (143000), 319 (sh, 43000), 380 (sh, 15000), 438 (3900), 488 (sh, 3700), 622 (sh, 400) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.57$  (m, 2 H), 4.50 (m, 2 H), 4.39 (m, 2 H), 4.26 (m, 2 H), 1.99 (quint,  $J = 7$  Hz, 4 H), 1.84 (quint.,  $J = 7$  Hz, 4 H), 1.75 (quint.,  $J = 7$  Hz, 4 H), 1.63 (quint.,  $J = 7$  Hz, 4 H), 1.07 (s, 9 H), 1.04 (s, 9 H), 1.03 (s, 9 H), 0.95 (s, 9 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 163.3$ , 163.2, 148.2, 147.4, 147.3 (two peaks), 146.8, 146.2, 146.0, 145.8, 145.7, 145.5, 145.3, 145.0 (two peaks), 144.7, 144.6, 144.4 (two peaks), 144.1, 143.6 (two peaks), 143.5, 142.5, 142.4, 141.3, 141.1, 140.1, 137.1, 136.4, 136.1, 135.6, 70.6, 67.7, 66.7 (two peaks), 63.2, 62.8, 50.3, 28.8, 28.5, 27.9 (three peaks), 27.8, 26.9, 25.5, 25.0, 21.4, 21.3, 21.2 ppm. MALDI-TOF-MS: 1515.38 (100%, [M+Na]<sup>+</sup>, calcd. for C<sub>98</sub>H<sub>68</sub>O<sub>12</sub>Si<sub>2</sub>Na: 1515.41), 1493.33 (81%, [M+H]<sup>+</sup>, calcd. for C<sub>98</sub>H<sub>69</sub>O<sub>12</sub>Si<sub>2</sub>: 1493.43), 1531.36 (26%, [M+K]<sup>+</sup>, calcd. for C<sub>98</sub>H<sub>68</sub>O<sub>12</sub>Si<sub>2</sub>K: 1531.39).

**Compounds II-22c & II-21c:** **II-22c & II-21c** were synthesized from **II-20c** (430 mg, 0.52 mmol), C<sub>60</sub> (374 mg, 0.52 mmol), I<sub>2</sub> (330 mg, 1.3 mmol) and DBU (0.36 mL, 2.3 mmol) in toluene (800 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5 to 75:25) yielded **II-22c** (95 mg, 0.06 mmol, 12%) as a red-brown glassy solid and **II-21c** (350 mg, 0.23 mmol, 44%) as a brown glassy solid. **II-22c:** IR (neat):  $\nu = 1744$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 249$  (48200), 308 (sh, 17200), 398 (3000), 409 (sh, 1800), 422 (1800), 480 (2100), 620 (sh, 300) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.73$ -4.53 (m, 4 H), 4.36 (m, 2 H), 4.25 (m, 2 H), 3.88 (m, 2 H), 3.83-3.66 (m, 5 H), 3.54 (m, 1 H), 1.77 (m, 8 H), 1.66 – 1.50 (m, 16 H), 1.00 (s, 9 H), 0.97 (s, 18 H), 0.90 (s, 9 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 164.3$ , 163.6, 163.0, 162.8, 149.0, 147.4, 147.2, 147.1, 146.6, 146.5, 146.3 (two peaks), 146.1, 146.0 (two peaks), 145.7, 145.6, 145.4 (two peaks), 145.1, 145.0, 144.9, 144.8, 144.7 (two peaks), 144.6 (two peaks), 144.3 (two peaks), 144.1, 143.9 (two peaks), 143.8, 143.7 (two peaks), 143.5, 143.4, 143.3, 143.2, 143.1 (two peaks), 142.9, 142.4, 142.0, 141.7, 141.4, 141.2, 140.0, 139.9, 139.7, 139.5, 138.7, 138.4, 71.9, 71.8, 71.7, 70.4, 67.8, 67.4, 66.9, 64.3, 64.0, 63.4 (two peaks), 53.8, 53.1, 33.3, 32.8, 32.3, 32.2, 29.2, 28.3, 28.2, 27.9 (several peaks), 27.7, 23.5, 23.2, 22.3, 22.2, 21.3 (two peaks), 21.1 (two peaks) ppm. MALDI-TOF-MS: 1571.82 ([M+Na]<sup>+</sup>, calcd. for C<sub>102</sub>H<sub>76</sub>O<sub>12</sub>Si<sub>2</sub>Na: 1571.48). **II-21c:** IR (neat):  $\nu = 1749$  (C=O), 1724 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 250$  (49300), 319 (sh, 15300), 399 (sh, 3400), 412 (2800), 424 (2300), 493 (2000), 574 (sh, 900), 628 (sh, 400), 686 (sh, 200) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.81$  (m, 2 H), 4.57 (m, 2 H), 4.40 (m, 4 H), 3.62-2.52 (m, 8 H), 1.90 (m, 2 H), 1.75 (m, 6 H), 1.50 – 1.34 (m, 16 H), 0.93 (s, 18 H), 0.92 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 164.8$ , 163.4, 147.5 (two peaks), 147.3, 147.2, 146.6, 146.4 (two peaks), 146.3, 145.5, 145.4, 145.1, 144.8, 144.4, 144.3, 144.0, 143.7, 143.6, 143.5, 142.7, 142.6, 142.0, 141.7, 141.1 (two peaks), 140.0, 139.6, 138.8, 71.7, 71.4, 68.2, 67.3, 64.2, 64.0, 53.4, 32.7, 32.5, 28.7, 27.9, 27.8, 26.9, 23.4, 21.1 (two peaks) ppm. MALDI-TOF-MS: 1571.80 ([M+Na]<sup>+</sup>, calcd. for C<sub>102</sub>H<sub>76</sub>O<sub>12</sub>Si<sub>2</sub>Na: 1571.48).

**Compounds II-22d & II-21d:** **II-22d & II-21d** were synthesized from **II-20d** (210 mg, 0.24 mmol), C<sub>60</sub> (170 mg, 0.24 mmol), I<sub>2</sub> (149 mg, 0.59 mmol) and DBU (0.16 mL, 1.1 mmol) in toluene (340

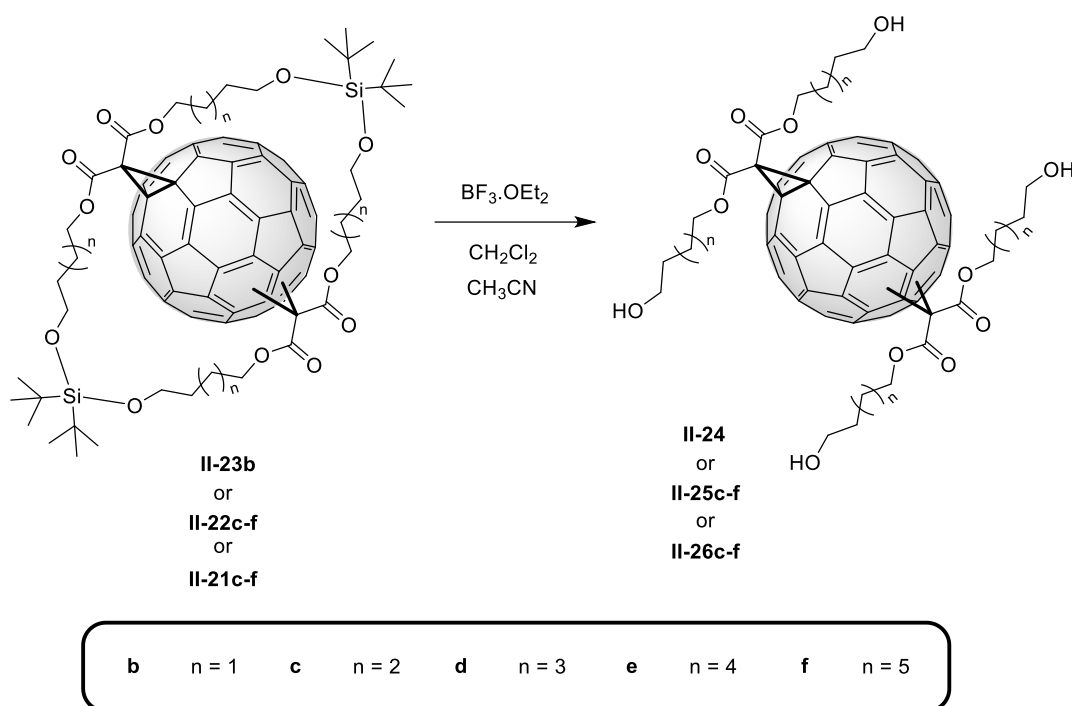
mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 2:3 to 3:2) yielded **II-22d** (102 mg, 0.06 mmol, 27%) as a red-brown glassy solid and **II-21d** (125 mg, 0.07 mmol, 33%) as a brown glassy solid. **II-22d**: IR (neat):  $\nu = 1745$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 253$  (137000), 309 (sh, 53200), 356 (sh, 21300), 397 (5000), 409 (sh, 3100), 420 (3000), 481 (3700), 618 (sh, 600) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.72$  (m, 1 H), 4.57 (m, 3 H), 4.33 (m, 2 H), 4.16 (m, 2 H), 3.80 (m, 7 H), 3.71 (m, 1 H), 1.77 (m, 8 H), 1.64 – 1.13 (m, 24 H), 1.00 (s, 9 H), 0.97 (s, 18 H), 0.91 (s, 9 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 163.7, 163.5, 163.2, 147.4, 147.2, 146.6, 146.4, 146.1, 145.7, 145.6, 145.4, 145.2, 144.8, 144.7, 144.6, 144.4, 144.1, 144.0, 143.9$  (two peaks), 143.5 (two peaks), 143.3, 142.5, 142.1, 141.6 (two peaks), 140.6, 139.6 (two peaks), 138.6 (two peaks), 71.8, 67.7, 67.3, 66.9 (two peaks), 64.2, 63.6, 63.4, 51.9, 33.2, 32.8, 32.7 (two peaks), 28.6, 28.2, 28.0, 27.9, 27.2, 26.9, 26.5, 26.1, 25.5, 25.2, 25.1 (two peaks), 25.0 (two peaks), 21.1 (three peaks) ppm. MALDI-TOF-MS: 1627.65 (100%, [M+Na]<sup>+</sup>, calcd. for C<sub>106</sub>H<sub>84</sub>O<sub>12</sub>Si<sub>2</sub>Na: 1627.54), 1643.59 (30%, [M+K]<sup>+</sup>, calcd. for C<sub>106</sub>H<sub>84</sub>O<sub>12</sub>Si<sub>2</sub>K: 1643.51), 1605.61 (12%, [M]<sup>+</sup>, calcd. for C<sub>106</sub>H<sub>85</sub>O<sub>12</sub>Si<sub>2</sub>: 1605.56). **II-21d**: IR (neat):  $\nu = 1749$  (C=O), 1727 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 248$  (172800), 322 (sh, 52000), 376 (sh, 17000), 399 (sh, 5000), 412 (5100), 424 (4300), 490 (3600), 570 (sh, 1800), 632 (sh, 700), 694 (sh, 260) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.78$  (m, 2 H), 4.52 (m, 2 H), 4.38 (m, 4 H), 3.64 (m, 8 H), 1.80 (m, 4 H), 1.69 (m, 4 H), 1.50 – 1.19 (m, 24 H), 0.93 (s, 18 H), 0.91 (s, 18 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 164.4, 163.8, 147.3, 147.2, 147.0, 146.9, 146.7$  (two peaks), 146.5, 146.4 (two peaks), 145.8, 145.4, 145.3, 144.7, 144.4, 144.3, 144.0, 143.6 (two peaks), 143.0, 142.6, 142.0, 141.7 (two peaks), 141.6, 140.2, 139.0, 138.2, 71.7, 71.6, 67.5, 63.8, 63.7, 53.1, 33.0, 32.9, 29.7, 28.8, 28.7, 27.6, 26.5, 26.3, 25.6, 25.4, 21.2, 21.1 ppm. MALDI-TOF-MS: 1627.56 ([M+Na]<sup>+</sup>, calcd. for C<sub>106</sub>H<sub>84</sub>O<sub>12</sub>Si<sub>2</sub>Na: 1627.54).

**Compounds II-22e & II-21e:** **II-22e & II-21e** were synthesized from **II-20e** (288 mg, 0.30 mmol), C<sub>60</sub> (220 mg, 0.30 mmol), I<sub>2</sub> (193 mg, 0.76 mmol) and DBU (0.21 mL, 1.4 mmol) in toluene (440 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 3:2 to 2:1) yielded **II-22e** (150 mg, 0.09 mmol, 30%) as a red-brown glassy solid and **II-21e** (102 mg, 0.06 mmol, 20%) as a brown glassy solid. **II-22e**: IR (neat):  $\nu = 1747$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 252$  (135000), 307 (sh, 52000), 356 (sh, 20000), 397 (5300), 408 (sh, 3300), 421 (3000), 482 (3700), 617 (sh, 600) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.68 - 4.46$  (m, 4 H), 4.35 – 4.19 (m, 4 H), 3.82 – 3.74 (m, 6 H), 3.67 (t, *J* = 7 Hz, 8 H), 1.76 (m, 8 H), 1.53 (m, 8 H), 1.49 – 1.13 (m, 24 H), 0.99 (s, 9 H), 0.98 (s, 9 H), 0.97 (s, 9 H), 0.93 (s, 9 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 163.9, 163.6, 163.5, 163.3, 148.6, 147.5, 147.3, 147.2, 146.5$  (two peaks), 146.4, 146.2, 146.1 (two peaks), 146.0, 145.6, 145.5, 145.2, 145.1 (two peaks), 145.0 (two peaks), 144.8, 144.7 (two peaks), 144.6 (three peaks), 144.5, 144.3 (two peaks), 144.1, 144.0, 143.8 (two peaks), 143.7 (three peaks), 143.5, 143.3 (two peaks), 143.0 (two peaks), 142.7, 142.3, 142.3, 142.2, 142.1, 141.9 (two peaks), 141.6, 141.5, 141.1, 140.8, 139.3, 138.6 (two peaks), 71.8, 70.4, 67.4, 67.3, 67.2, 63.9 (two peaks), 63.7, 63.6, 53.8, 51.9, 33.1, 32.8, 32.8, 32.7, 32.3, 29.5, 29.0, 28.7, 28.6 (two peaks), 28.4 (two peaks), 28.2, 27.9 (two peaks), 26.8, 26.0, 25.9, 25.8, 25.7, 25.5, 25.4, 21.2, 21.1 (two peaks) ppm. MALDI-TOF-MS: 1683.61 (100%, [M+Na]<sup>+</sup>, calcd. for C<sub>110</sub>H<sub>93</sub>O<sub>12</sub>Si<sub>2</sub>Na:



1683.60), 1663.64 (77%, [M]<sup>+</sup>, calcd. for C<sub>110</sub>H<sub>93</sub>O<sub>12</sub>Si<sub>2</sub>: 1662.62). **II-21e**: IR (neat):  $\nu = 1742$  (C=O), 1728 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 254$  (134500), 330 (sh, 35700), 399 (sh, 5300), 409 (4200), 424 (3300), 488 (2800), 583 (sh, 1100), 627 (sh, 500), 688 (sh, 200) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.77$  (m, 2 H), 4.43 (m, 6 H), 3.66 (m, 8 H), 1.82 (m, 4 H), 1.70 (m, 4 H), 1.50 – 1.19 (m, 32 H), 0.93 (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 163.9$  (two peaks), 147.2, 147.0, 146.8, 146.6 (two peaks), 146.5, 146.4, 146.3 (two peaks), 146.0, 145.5, 145.4, 144.6, 144.3, 144.2, 143.9, 143.6, 143.4, 142.9, 142.5, 142.2 (two peaks), 141.9, 141.6, 140.3, 138.5, 137.8, 71.9, 71.6, 67.5, 67.3, 63.8, 63.7, 53.0, 32.8, 32.7, 29.0, 28.8, 28.7, 28.6, 27.9 (two peaks), 26.4, 26.0, 25.9, 25.6, 21.1 (two peaks) ppm. MALDI-TOF-MS: 1661.66 (100%, [M]<sup>+</sup>, calcd. for C<sub>110</sub>H<sub>93</sub>O<sub>12</sub>Si<sub>2</sub>: 1662.62), 1683.66 (52%, [M+Na]<sup>+</sup>, calcd. for C<sub>110</sub>H<sub>93</sub>O<sub>12</sub>Si<sub>2</sub>Na: 1683.60).

**Compound II-22f & II-38f: II-22f & II-21f** were synthesized from **II-20f** (280 mg, 0.28 mmol), C<sub>60</sub> (201 mg, 0.28 mmol), I<sub>2</sub> (178 mg, 0.70 mmol) and DBU (0.19 mL, 1.3 mmol) in toluene (400 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 3:2 to 2:1) yielded **II-22f** (102 mg, 0.06 mmol, 21%) as a red-brown glassy solid and **II-21f** (79 mg, 0.05 mmol, 16%) as a brown glassy solid. **II-22f**: IR (neat):  $\nu = 1744$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 256$  (112000), 313 (sh, 42000), 356 (sh, 17000), 393 (4600), 407 (sh, 3000), 422 (2700), 480 (3200), 628 (sh, 300) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.63 - 4.23$  (m, 8 H), 3.79 (m, 4 H), 3.69 (t,  $J = 7$  Hz, 4 H), 1.75 (m, 8 H), 1.56 (m, 8 H), 1.48 – 1.13 (m, 40 H), 1.00 (s, 9 H), 0.99 (s, 9 H), 0.96 (s, 9 H), 0.94 (s, 9 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 163.9$ , 163.6, 163.4, 148.5, 147.5, 147.3, 147.2, 146.4 (two peaks), 146.2, 146.1 (two peaks), 146.0, 145.6, 145.5, 145.3, 145.2 (two peaks), 145.0, 144.9, 144.8, 144.7 (two peaks), 144.6, 144.5, 144.4, 144.2, 144.1, 144.0 (two peaks), 143.8, 143.7, 143.5, 143.3 (two peaks), 142.9 (two peaks), 142.8, 142.6, 142.3, 142.0, 141.9, 141.6, 141.5, 141.2, 140.8, 138.9 (two peaks), 138.6 (two peaks), 71.9, 71.6, 70.4, 67.4, 67.3, 67.2, 63.9, 63.8, 63.5, 53.8, 51.9, 33.1, 32.9, 32.8, 29.7, 29.4, 29.3 (two peaks), 29.2 (two peaks), 28.9 (two peaks), 28.8, 28.7, 28.4 (two peaks), 27.9, 26.5, 26.3, 26.0, 25.7, 25.6 (two peaks), 25.0 (two peaks), 21.1 (three peaks) ppm. MALDI-TOF-MS: 1718.00 ([M+H]<sup>+</sup>, calcd. for C<sub>114</sub>H<sub>101</sub>O<sub>12</sub>Si<sub>2</sub>: 1718.69). **II-21f**: IR (neat):  $\nu = 1751$  (C=O), 1732 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 249$  (132000), 318 (sh, 41000), 397 (sh, 5300), 410 (4200), 424 (3300), 487 (2900), 573 (sh, 1300), 627 (sh, 500), 688 (sh, 100) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.73$  (m, 2 H), 4.41 (m, 6 H), 3.70 (m, 8 H), 1.82 (m, 4 H), 1.69 (m, 4 H), 1.50 – 1.19 (m, 40 H), 0.95 (s, 36 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 163.9$ , 163.8, 147.2, 147.0, 146.8, 146.6, 146.5, 146.3 (two peaks), 146.1, 145.5, 145.4, 144.6, 144.4, 144.1, 143.8, 143.5, 143.4, 143.0, 142.5, 142.2, 141.9, 141.6, 140.4, 138.6, 138.4, 137.7, 71.8, 71.6, 67.5, 67.2, 63.8, 52.9, 32.9 (two peaks), 29.4 (two peaks), 29.1, 29.0, 28.7, 28.6, 27.9 (two peaks), 26.2, 26.0, 25.9, 25.7, 21.1 (two peaks) ppm. MALDI-TOF-MS: 1719.83 ([M+H]<sup>+</sup>, calcd. for C<sub>114</sub>H<sub>101</sub>O<sub>12</sub>Si<sub>2</sub>: 1718.69).



**General Procedure for the desilylation of bis-adducts II-23b or II-22c-f or II-21c-f :**  $\text{BF}_3 \cdot \text{OEt}_2$  (10-15 eq.) was added to a solution of the appropriate bis-adducts (1 eq.) in  $\text{CH}_2\text{Cl}_2/\text{CH}_3\text{CN}$  (2:1). The resulting mixture was stirred overnight at room temperature, then filtered on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 95:5) and concentrated. Column chromatography on  $\text{SiO}_2$  gave **II-24 or II-25c-f and II-26c-f**.

**Compound II-24:** **II-24** was synthesized from **II-23b** (77 mg, 52  $\mu\text{mol}$ ) and  $\text{BF}_3 \cdot \text{OEt}_2$  (0.1 mL, 790  $\mu\text{mol}$ ) in  $\text{CH}_2\text{Cl}_2/\text{CH}_3\text{CN}$  (3/1.5 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 97:3 to 92:8) yielded **II-24** (52 mg, 43  $\mu\text{mol}$ , 88%) as a brown glassy solid. IR (neat):  $\nu = 3340$  (br, OH), 1741 (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 257$  (150000), 320 (sh, 47000), 379 (sh, 16000), 437 (4100), 490 (sh, 2900), 624 (sh, 500) nm.  $^1\text{H-NMR}$  (400 MHz,  $\text{CDCl}_3$ ):  $\delta = 4.40$  (m, 8 H), 3.68 (m, 8 H), 1.96 – 1.62 (m, 16 H) ppm.  $^{13}\text{C-NMR}$  (100 MHz,  $\text{CDCl}_3$ ):  $\delta = 163.4$ , 163.1, 148.4, 147.4 (two peaks), 147.3, 146.2, 145.9 (two peaks), 145.8, 145.7, 145.4, 145.3, 145.1, 145.0, 144.7, 144.4 (two peaks), 144.2, 143.7, 143.6, 143.4, 142.5, 142.4, 141.3, 141.0, 139.7, 137.8, 137.1, 136.7, 135.2, 70.5, 67.7, 67.1, 62.2, 62.1, 49.6, 29.2, 29.1, 25.1, 25.0 ppm. MALDI-TOF-MS: 1212.44 ( $[\text{M}]^+$ , calcd. for  $\text{C}_{82}\text{H}_{36}\text{O}_{12}$ : 1212.22).

**Compound II-25c:** **II-25c** was synthesized from **II-22c** (82 mg, 53  $\mu\text{mol}$ ) and  $\text{BF}_3 \cdot \text{OEt}_2$  (0.1 mL, 790  $\mu\text{mol}$ ) in  $\text{CH}_2\text{Cl}_2/\text{CH}_3\text{CN}$  (3/1.5 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{THF}$ , 3:1 to  $\text{THF}/\text{MeOH}$ , 95:5) yielded **II-25c** (57 mg, 45  $\mu\text{mol}$ , 85%) as a red-brown glassy solid. IR (neat):  $\nu = 3339$  (br, OH), 1739 (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 240$  (29400), 308 (sh, 11600), 398 (1900), 410 (sh, 1200), 422 (1200), 481 (1400), 620 (sh, 200) nm.  $^1\text{H-NMR}$  (400 MHz,  $\text{CDCl}_3$ ):  $\delta = 4.43$  (m, 8 H), 3.64 (m, 8 H), 1.85 (m, 12 H), 1.60 (m, 8 H), 1.51 (m, 8 H) ppm.  $^{13}\text{C-NMR}$  (100 MHz,  $\text{CDCl}_3$ ):  $\delta = 163.7$ , 163.6, 163.5, 147.5, 147.2, 146.4, 146.1 (two peaks), 145.6,

145.3, 145.1, 145.0, 144.7 (two peaks), 144.6, 144.3 (two peaks), 144.2, 144.0, 143.7 (two peaks), 143.4, 143.1, 142.9, 142.8, 142.3, 141.9, 141.6 (two peaks), 138.9, 138.7, 71.7, 71.6, 70.4, 67.3, 67.2, 62.5, 53.7, 53.44, 51.5, 32.2, 32.1, 28.3, 22.3 (two peaks) ppm. MALDI-TOF-MS: 1286.47 ([M]<sup>+</sup>, calcd for C<sub>86</sub>H<sub>44</sub>O<sub>12</sub>: 1268.28).

**Compound II-25d:** **II-25d** was synthesized from **II-22d** (82 mg, 51 μmol) and BF<sub>3</sub>.OEt<sub>2</sub> (0.1 mL, 790 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/CH<sub>3</sub>CN (3/1.5 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 92:8) yielded **II-25d** (54 mg, 41 μmol, 80%): red-brown glassy solid. IR (neat): ν = 3338 (br, OH), 1741 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>): λ<sub>max</sub>(ε) = 251 (133600), 310 (sh, 50600), 358 (sh, 19100), 396 (sh, 5000), 406 (3100), 421 (2900), 482 (3600), 620 (sh, 500) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 4.42 (m, 8 H), 3.64 (m, 8 H), 1.81 (m, 8 H), 1.58 (m, 8 H), 1.51 – 1.38 (m, 12 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 163.7, 163.6 (two peaks), 147.6, 147.3, 146.5, 146.2, 146.1, 145.6, 145.4, 145.2, 145.1, 144.7 (two peaks), 144.6, 144.4 (two peaks), 144.2, 144.1, 143.8, 143.7, 143.5, 143.2, 143.0, 142.4, 141.9, 141.6 (two peaks), 138.9, 138.7, 71.7, 71.6, 70.4, 67.3, 62.7, 53.7, 51.5, 32.6 (two peaks), 28.5 (two peaks), 28.4, 25.8 (two peaks), 25.7, 25.4 (two peaks), 25.3 ppm. MALDI-TOF-MS: 1324.45 (100%, [M]<sup>+</sup>, calcd for C<sub>90</sub>H<sub>52</sub>O<sub>12</sub>: 1324.34), 1347.35 (15%, [M+Na]<sup>+</sup>, calcd. for C<sub>90</sub>H<sub>52</sub>O<sub>12</sub>Na: 1347.34), 1363.26 (20%, [M+K]<sup>+</sup>, calcd. for C<sub>90</sub>H<sub>52</sub>O<sub>12</sub>K: 1363.31).

**Compound II-25e:** **II-25e** was synthesized from **II-22e** (95 mg, 57 μmol) and BF<sub>3</sub>.OEt<sub>2</sub> (0.1 mL, 790 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/CH<sub>3</sub>CN (3/1.5 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 93:7) yielded **II-25e** (55 mg, 40 μmol, 70%) as a red-brown glassy solid. IR (neat): ν = 3316 (br, OH), 1741 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>): λ<sub>max</sub>(ε) = 251 (151000), 309 (sh, 58700), 356 (sh, 23300), 396 (sh, 5000), 398 (5300), 409 (sh, 3400), 422 (3100), 480 (sh, 3800), 626 (sh, 500) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 4.40 (m, 8 H), 3.64 (m, 8 H), 1.79 (m, 8 H), 1.58 (m, 16 H), 1.48 – 1.34 (m, 24 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 163.6, 163.5, 147.6, 147.2, 146.4, 146.1, 146.0, 145.5, 145.3, 145.1, 145.0, 144.7 (two peaks), 144.6, 144.4 (two peaks), 144.1, 144.0, 143.7 (two peaks), 143.4, 143.3, 143.1, 142.9, 142.3, 141.9, 141.6, 141.5, 138.7, 138.6, 71.7, 70.5, 67.3, 62.8, 53.7, 51.5, 32.6, 28.9 (two peaks), 28.5, 25.8 (two peaks), 25.9, 25.8, 25.7 ppm. MALDI-TOF-MS: 1380.36 ([M]<sup>+</sup>, calcd for C<sub>94</sub>H<sub>60</sub>O<sub>12</sub>: 1380.41).

**Compound II-25f:** **II-25f** was synthesized from **II-22f** (73 mg, 43 μmol) and BF<sub>3</sub>.OEt<sub>2</sub> (0.1 mL, 790 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/CH<sub>3</sub>CN (3/1.5 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 94:6) yielded **II-25f** (50 mg, 35 μmol, 82%) as a red-brown glassy solid. IR (neat): ν = 3330 (br, OH), 1741 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>): λ<sub>max</sub>(ε) = 257 (82000), 315 (sh, 31500), 398 (sh, 3500), 410 (sh, 2200), 423 (2000), 483 (sh, 2600), 636 (sh, 190) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 4.40 (m, 8 H), 3.64 (m, 8 H), 1.79 (m, 8 H), 1.58 (m, 16 H), 1.45 – 1.34 (m, 32 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 163.6 (two peaks), 163.5, 147.6, 147.2, 146.4, 146.1, 146.0, 145.5, 145.3, 145.1, 145.0, 144.7, 144.6, 144.5, 144.4, 144.0 (two peaks), 143.8, 143.7, 143.4, 143.3, 143.2, 142.9, 142.3, 141.8, 141.6, 141.5, 138.6, 71.7, 70.4, 67.3, 62.9, 53.7, 51.5, 32.7, 29.3 (two peaks), 29.2 (two peaks), 29.1, 28.5 (two peaks), 28.4, 25.9 (two peaks), 25.8, 25.7 (two peaks), 25.6 ppm. MALDI-TOF-MS: 1436.26 ([M]<sup>+</sup>, calcd for C<sub>98</sub>H<sub>68</sub>O<sub>12</sub>: 1436.47).

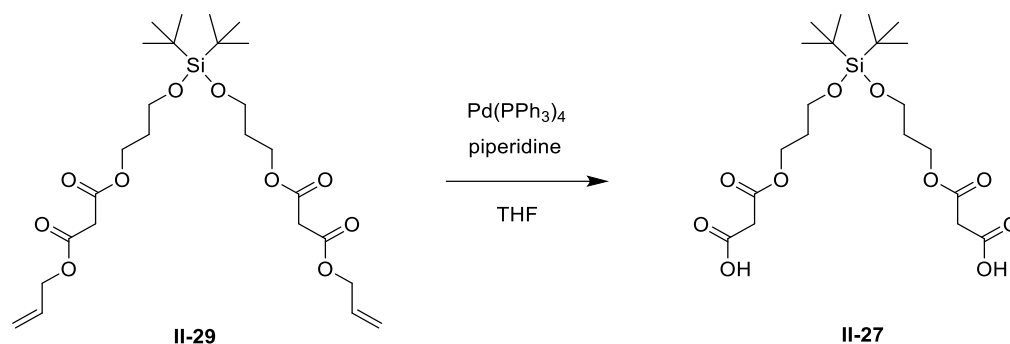
**Compound II-26c:** **II-26c** was synthesized from **II-21c** (118 mg, 76  $\mu$ mol) and BF<sub>3</sub>.OEt<sub>2</sub> (0.2 mL, 1.5 mmol) in CH<sub>2</sub>Cl<sub>2</sub>/CH<sub>3</sub>CN (3/1.5 mL). Column chromatography on SiO<sub>2</sub> (THF to THF/MeOH, 95:5) yielded **II-26c** (78 mg, 61  $\mu$ mol, 81%) as a brown glassy solid. IR (neat):  $\nu$  = 3326 (br, OH), 1737 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon)$  = 248 (60200), 319 (sh, 27300), 400 (sh, 3000), 412 (2500), 424 (2100), 490 (1700), 576 (sh, 800), 633 (sh, 330), 689 (sh, 100) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.52 (t, *J* = 7 Hz, 4 H), 4.42 (2t, *J* = 7 Hz, 4 H), 3.68 (t, *J* = 7 Hz, 4 H), 3.62 (t, *J* = 7 Hz, 4 H), 1.90 (m, 4 H), 1.80 (m, 4 H), 1.75 (OH, 4 H), 1.65 – 1.47 (m, 16 H). <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>) :  $\delta$  = 163.6 (two peaks), 147.2, 147.0, 146.6, 146.5, 146.4 (two peaks), 146.1, 146.0, 145.6, 145.5, 145.3, 144.6, 144.4, 144.2, 143.8, 143.5, 143.4, 143.0, 142.5, 142.2, 142.1, 141.9, 141.6, 140.3, 139.1, 138.4, 71.8, 71.3, 67.3, 67.2, 62.5 (two peaks), 51.8, 32.2, 32.1, 28.4, 28.3, 22.4, 22.3 ppm. MALDI-TOF-MS: 1286.35 (100%, [M]<sup>+</sup>, calcd for C<sub>86</sub>H<sub>44</sub>O<sub>12</sub>: 1268.28), 1291.34 (27%, [M+Na]<sup>+</sup>, calcd. C<sub>86</sub>H<sub>44</sub>O<sub>12</sub>: 1291.27).

**Compound II-26d:** **II-26d** was synthesized from **II-21d** (96 mg, 60  $\mu$ mol) and BF<sub>3</sub>.OEt<sub>2</sub> (0.1 mL, 790  $\mu$ mol) in CH<sub>2</sub>Cl<sub>2</sub>/CH<sub>3</sub>CN (3/1.5 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 92:8) yielded **II-26d** (70 mg, 53  $\mu$ mol, 89%) as a brown glassy solid. IR (neat):  $\nu$  = 3325 (br, OH), 1741 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon)$  = 248 (132500), 319 (sh, 40700), 381 (sh, 10900), 400 (sh, 4900), 411 (4100), 422 (3300), 490 (2800), 576 (sh, 1300), 632 (sh, 500), 693 (sh, 180) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.51 (t, *J* = 7 Hz, 4 H), 4.40 (t, *J* = 7 Hz, 4 H), 3.66 (t, *J* = 6 Hz, 4 H), 3.62 (t, *J* = 6 Hz, 4 H), 1.91 – 1.73 (m, 12 H), 1.65 – 1.38 (m, 20 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>) :  $\delta$  = 163.6 (two peaks), 147.2, 147.0, 146.6, 146.5 (two peaks), 146.4, 146.2, 146.1, 145.6, 145.5, 145.4, 144.7, 144.4, 144.2, 143.9, 143.6, 143.4, 143.1, 142.6, 142.2, 142.1, 142.0, 141.7, 140.4, 139.2, 138.5, 71.8, 71.3, 67.4, 67.3, 62.8 (two peaks), 51.8, 32.6 (two peaks), 28.6, 28.5, 25.9, 25.8, 25.5, 25.4 ppm. MALDI-TOF-MS: 1324.38 ([M]<sup>+</sup>, calcd for C<sub>90</sub>H<sub>52</sub>O<sub>12</sub>: 1324.35).

**Compound II-26e:** **II-26e** was synthesized from **II-21e** (70 mg, 42  $\mu$ mol) and BF<sub>3</sub>.OEt<sub>2</sub> (0.1 mL, 790  $\mu$ mol) in CH<sub>2</sub>Cl<sub>2</sub>/CH<sub>3</sub>CN (3/1.5 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) yielded **II-26e** (36 mg, 26  $\mu$ mol, 66%) as a brown glassy solid. IR (neat):  $\nu$  = 3336 (br, OH), 1743 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon)$  = 248 (137000), 317 (sh, 43000), 397 (sh, 5400), 411 (4400), 422 (3600), 492 (3000), 581 (sh, 1200), 635 (sh, 500), 689 (sh, 160) nm. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.51 (t, *J* = 7 Hz, 4 H), 4.40 (t, *J* = 7 Hz, 4 H), 3.65 (t, *J* = 7 Hz, 4 H), 3.62 (t, *J* = 7 Hz, 4 H), 1.87 (m, 4 H), 1.77 (m, 4 H), 1.56 (m, 16 H), 1.43 – 1.34 (m, 20 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>) :  $\delta$  = 163.6 (two peaks), 147.2, 146.9, 146.5 (two peaks), 146.4, 146.1, 146.0, 145.6, 145.5, 145.3, 144.6, 144.4, 144.1, 143.8, 143.5 (two peaks), 143.4, 143.0, 142.5, 142.2, 142.1, 141.9, 140.6, 140.3, 139.1, 138.4, 71.8, 71.4, 67.4, 67.3, 62.8 (two peaks), 51.9, 32.6 (two peaks), 29.0 (two peaks), 28.6, 28.5, 25.9 (two peaks), 25.8, 25.7 ppm. MALDI-TOF-MS: 1380.40 ([M]<sup>+</sup>, calcd for C<sub>94</sub>H<sub>60</sub>O<sub>12</sub>: 1380.41).

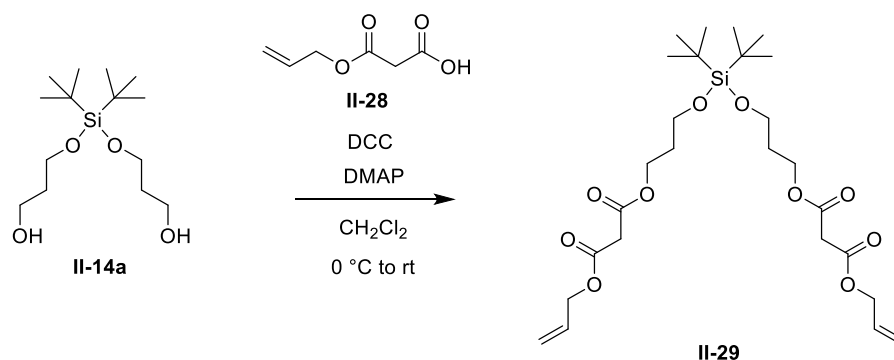
**Compound II-26f:** **II-26f** was synthesized from **II-21f** (60 mg, 35  $\mu$ mol) and BF<sub>3</sub>.OEt<sub>2</sub> (0.1 mL, 790  $\mu$ mol) in CH<sub>2</sub>Cl<sub>2</sub>/CH<sub>3</sub>CN (3/1.5 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 94:6) yielded **II-26f** (28 mg, 19  $\mu$ mol, 56%) as a brown glassy solid. IR (neat):  $\nu$  = 3330 (br, OH), 1746 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon)$  = 248 (126000), 319 (sh, 41000), 398 (sh, 5000), 410 (4200), 423 (3400), 488 (sh, 3000), 579 (sh, 1200), 632 (sh, 500), 689 (sh, 150) nm. <sup>1</sup>H-NMR

(400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.51 (t,  $J$  = 7 Hz, 4 H), 4.40 (t,  $J$  = 7 Hz, 4 H), 3.64 (t,  $J$  = 7 Hz, 4 H), 3.63 (t,  $J$  = 7 Hz, 4 H), 1.86 (m, 4 H), 1.77 (m, 4 H), 1.56 (m, 16 H), 1.43 – 1.31 (m, 28 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta$  = 163.6 (two peaks), 147.2, 146.9, 146.5, 146.4 (two peaks), 146.2, 146.0, 145.6, 145.5, 145.3, 144.6, 144.4, 144.1, 143.8, 143.5 (two peaks), 143.4, 143.0, 142.5, 142.2, 142.1, 141.9, 141.6, 140.3, 139.1, 138.4, 71.8, 71.4, 67.4, 67.3, 62.9 (two peaks), 51.9, 32.7 (two peaks), 29.4 (two peaks), 29.2 (two peaks), 28.6, 28.5, 26.0, 25.9, 25.7 (two peaks) ppm. MALDI-TOF-MS: 1436.46 ([M]<sup>+</sup>, calcd for C<sub>98</sub>H<sub>68</sub>O<sub>12</sub>: 1436.47).

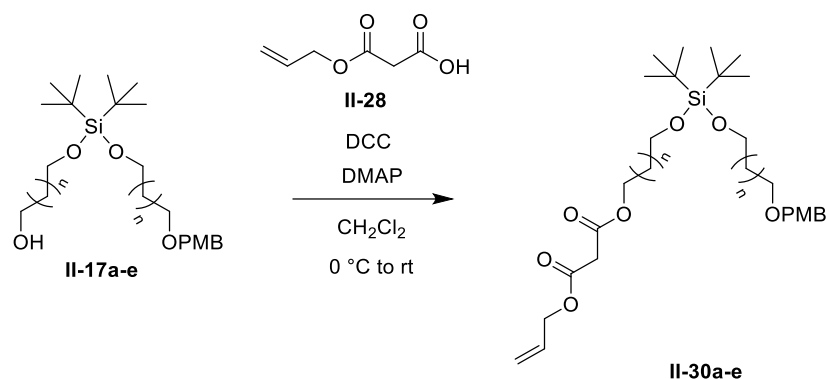


**Compound II-27:** Piperidine (5.8 mL, 58 mmol) and Pd(PPh<sub>3</sub>)<sub>4</sub> (0.54 g, 0.47 mmol) were added to a solution of **II-29** (1.28 g, 2.3 mmol) in THF (10 mL). After 4 h, the solvent was removed under vacuum. The crude was dissolved in CH<sub>2</sub>Cl<sub>2</sub> then the solution was acidified with HCl 2M (10 mL). The organic layer was washed with water and the solvent was removed under vacuum to give **II-27**.

**Compound II-28:** **II-28** was prepared as described in Tetrahedron Letters, Vol. 38, No. 44, pp. 7737-7740, 1997.



**Compound II-29:** DCC (1.97 g, 9.55 mmol) was added to a solution of **II-14a** (0.93 g, 3.2 mmol), 10 (0.98 g, 6.8 mmol) and DMAP (0.39 g, 3.2 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (40 mL) at 0°C. The resulting mixture was stirred overnight at room temperature, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/Ethyl acetate, 9:1) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2) yielded **II-29** (1.27 g, 2.3 mmol, 74%) as a colorless oil.



**General procedure for the synthesis of compounds II-30a-e:** Dicyclohexylcarbodiimide (DCC, 2 eq) was added to a solution of **II-17a-e** (1 eq), **II-28** (1 eq) and 4-dimethylaminopyridine (DMAP, 1 eq) in CH<sub>2</sub>Cl<sub>2</sub> at 0 °C under argon. After 12 h the crude was filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> gave **II-30a-e**.

**Compound II-30a:** **II-30a** was synthesized from DCC (2.22 g, 10.8 mmol), **II-17a** (2.22 g, 5.38 mmol), **II-28** (0.853 g, 5.92 mmol) and DMAP (0.657 g, 5.38 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (20 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) gave **II-30a** (2.10 g, 73%) as a colorless oil. IR (neat):  $\nu = 1753$  (C=O), 1736 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.25$  (d,  $J = 8.5$  Hz, 2 H), 6.87 (d,  $J = 8.5$  Hz, 2 H), 5.90 (m, 1 H), 5.30 (m, 2 H), 4.64 (td,  $J = 5.5$  Hz, 1.5 Hz, 2 H), 4.43 (s, 2 H), 4.29 (t,  $J = 6.5$  Hz, 2 H), 3.92 (t,  $J = 6$  Hz, 2 H), 3.88 (t,  $J = 6$  Hz, 2 H), 3.80 (s, 3 H), 3.58 (t,  $J = 6.5$  Hz, 2 H), 3.39 (s, 2 H), 1.86 (m, 4 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C NMR (75 MHz, CDCl<sub>3</sub>):  $\delta = 166.5, 166.3, 159.2, 131.6, 130.8, 129.3, 188.8, 113.8, 72.8, 66.7, 66.1, 62.7, 60.8, 60.1, 55.3, 41.6, 33.2, 31.8, 27.9, 21.2$  ppm.

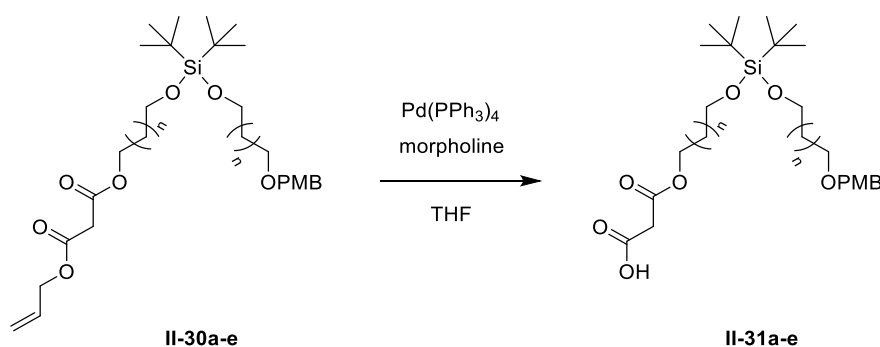
**Compound II-30b:** **II-30b** was synthesized from DCC (2.06 g, 10.0 mmol), **II-17b** (2.21 g, 5.02 mmol), **II-28** (0.796 g, 5.52 mmol) and DMAP (0.610 g, 5.02 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (40 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) gave **II-30b** (2.04 g, 71%) as a colorless oil. IR (neat):  $\nu = 1754$  (C=O), 1737 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.26$  (d,  $J = 8.5$  Hz, 2 H), 6.87 (d,  $J = 8.5$  Hz, 2 H), 5.91 (ddt,  $J = 17$  Hz, 10.5 Hz, 6 Hz, 5.5 Hz, 1 H), 5.29 (m, 2 H), 4.65 (m, 2 H), 4.44 (s, 2 H), 4.18 (t,  $J = 6.5$  Hz, 2 H), 3.82 (t,  $J = 6$  Hz, 2 H), 3.80 (s, 3 H), 3.48 (t,  $J = 6.5$  Hz, 2 H), 3.40 (s, 2 H), 1.68 (m, 8 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.6, 166.4, 159.3, 131.7, 130.9, 129.4, 118.9, 113.9, 72.6, 70.1, 66.2, 65.7, 63.8, 63.2, 55.4, 41.7, 29.8, 29.3, 28.0, 26.4, 25.3, 21.3$  ppm.

**Compound II-30c:** **II-30c** was synthesized from DCC (2.08 g, 10.1 mmol), **II-17c** (2.36 g, 5.04 mmol), **II-28** (0.80 g, 5.54 mmol) and DMAP (0.62 g, 5.04 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (20 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane/MeOH, 49:49:2) gave **II-30c** (3.01 g, 100%) as a light yellow oil. IR (neat):  $\nu = 1754$  (C=O), 1737 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.26$  (d,  $J = 8.5$  Hz, 2 H), 8.87 (d,  $J = 8.5$  Hz, 2 H), 5.90 (m, 1 H), 5.30 (m, 2 H), 4.65 (m, 2 H), 4.43 (s, 2 H), 4.15 (t,  $J = 6.5$  Hz, 2 H), 3.80 (m, 7 H), 3.45 (t,  $J = 6.5$  Hz, 2 H), 3.39 (s, 2 H), 1.59 (m, 8 H), 1.44 (m, 4 H), 0.99 (s, 18 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.7, 166.4, 159.3, 131.7,$

131.0, 129.3, 118.9, 113.9, 72.7, 70.3, 66.2, 65.8, 63.9, 63.6, 55.4, 41.7, 32.9, 32.5, 29.7, 28.4, 28.0, 22.7, 22.3, 21.3 ppm.

**Compound II-30d:** **II-30d** was synthesized from DCC (1.25 g, 6.1 mmol), **II-17d** (1.51 g, 3.03 mmol), **II-28** (0.44 g, 3.03 mmol) and DMAP (0.37 g, 3.0 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (8 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 7:3) gave **II-30d** (1.81 g, 96%) as a light yellow oil. IR (neat):  $\nu = 1754$  (C=O),  $1737$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.26$  (d,  $J = 8.5$  Hz, 2 H),  $6.87$  (d,  $J = 8.5$  Hz, 2 H),  $5.91$  (m, 1 H),  $5.30$  (m, 2 H),  $4.64$  (m, 2 H),  $4.43$  (s, 2 H),  $4.14$  (t,  $J = 6.5$  Hz, 2 H),  $3.80$  (m, 7 H),  $3.44$  (t,  $J = 6.5$  Hz, 2 H),  $3.40$  (s, 2 H),  $1.61$  (m, 8 H),  $1.38$  (m, 8 H),  $0.99$  (s, 18 H) ppm. <sup>13</sup>C NMR (75 MHz, CDCl<sub>3</sub>):  $\delta = 166.7, 166.4, 159.2, 131.7, 130.9, 129.3, 118.9, 113.9, 72.7, 70.3, 66.2, 65.8, 63.9, 63.7, 55.4, 41.7, 33.0, 32.9, 30.0, 28.6, 28.0, 26.2, 25.9, 25.8, 25.6, 21.3$  ppm.

**Compound II-30e:** **II-30e** was synthesized from DCC (1.63 g, 7.90 mmol), **II-17e** (2.07 g, 3.93 mmol), **II-28** (0.62 g, 4.33 mmol) and DMAP (0.48 g, 3.9 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (15 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 9:1) gave **II-30e** as a light yellow oil. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.26$  (d,  $J = 8.5$  Hz, 2 H),  $6.87$  (d,  $J = 8.5$  Hz, 2 H),  $5.91$  (m, 1 H),  $5.30$  (m, 2 H),  $4.65$  (d,  $J = 5.5$  Hz, 2 H),  $4.43$  (s, 2 H),  $4.14$  (t,  $J = 6.5$  Hz, 2 H),  $3.80$  (s, 3 H),  $3.79$  (t,  $J = 6.5$  Hz, 4 H),  $3.43$  (t,  $J = 6.5$  Hz, 2 H),  $3.40$  (s, 2 H),  $1.58$  (m, 8 H),  $1.35$  (m, 12 H),  $0.99$  (s, 18 H) ppm.



**General procedure for the synthesis of compounds II-31a-e:** Morpholine (5 eq) and Pd(PPh<sub>3</sub>)<sub>4</sub> (0.05 eq) were added to a solution of **II-30a-e** (1 eq) in dry THF under argon. After 4 h, the solvent was removed under vacuum. The crude was dissolved in CH<sub>2</sub>Cl<sub>2</sub> then the solution was acidified with HCl 2M (10 mL). The organic layer was washed with water and the solvent was removed under vacuum to give **II-31a-e**. The crude was used without purification.

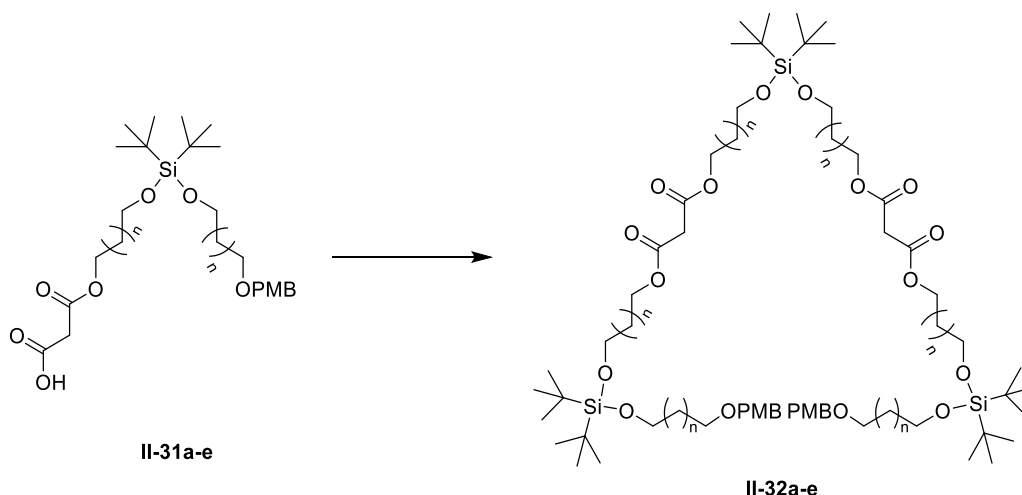
**Compound II-31a:** **II-31a** was synthesized from morpholine (2.0 mL, 22.9 mmol), Pd(PPh<sub>3</sub>)<sub>4</sub> (0.21 g, 0.18 mmol) and **II-430a** (0.49 g, 0.92 mmol) in dry THF (5 mL). **II-30a** was isolated as a yellow oil. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.26$  (d,  $J = 8.5$  Hz, 2 H),  $6.87$  (d,  $J = 8.5$  Hz, 2 H),  $4.49$  (s, 2 H),  $4.30$  (t,  $J = 6$  Hz, 2 H),  $3.90$  (m, 4 H),  $3.80$  (s, 3 H),  $3.61$  (t,  $J = 6$  Hz, 2 H),  $3.37$  (s, 2 H),  $1.87$  (m, 4 H),  $0.98$  (s, 18 H) ppm.

**Compound II-31b:** **II-31b** was synthesized from morpholine (2.9 mL, 34.0 mmol), Pd(PPh<sub>3</sub>)<sub>4</sub> (0.20 g, 0.17 mmol) and **II-30b** (2.04 g, 3.42 mmol) in dry THF (20 mL).

**Compound II-31c:** II-31c was synthesized from morpholine (6.61 mL, 75.9 mmol), Pd(PPh<sub>3</sub>)<sub>4</sub> (0.585 g, 0.506 mmol) and II-30c (3.01 g, 5.06 mmol) in dry THF (20 mL).

**Compound II-31d:** II-31d was synthesized from morpholine (1.3 mL, 14.5 mmol), Pd(PPh<sub>3</sub>)<sub>4</sub> (0.17 g, 0.145 mmol) and II-30d (1.81 g, 2.91 mmol) in dry THF (10 mL). <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.26 (d, J = 8.5 Hz, 2H), 6.87 (d, J = 8.5 Hz, 2H), 4.45 (s, 2H), 4.16 (t, J = 6.5 Hz, 2H), 3.80 (m, 7H), 3.45 (t, J = 6.5 Hz, 2H), 3.39 (s, 2H), 1.59 (m, 8H), 1.38 (m, 8H), 0.99 (s, 18H) ppm.

**Compound II-31e:** II-31e was synthesized from morpholine (4.3 mL, 48.8 mmol), Pd(PPh<sub>3</sub>)<sub>4</sub> (0.45 g, 0.39 mmol) and II-30e in dry THF (20 mL).



**General procedure for the synthesis of compounds II-32a-e:** Dicyclohexylcarbodiimide (DCC, 4.5 eq) was added to a solution of II-14a-e (1 eq), II-31a-e and 4-dimethylaminopyridine (DMAP, 2 eq) in CH<sub>2</sub>Cl<sub>2</sub> at 0 °C under argon. After 12 h the crude was filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>). Column chromatography (SiO<sub>2</sub>) gave II-32a-e.

**Compound II-32a:** II-32a was synthesized from DCC (0.28 g, 1.37 mmol), II-14a (0.13 g, 0.458 mmol), II-31a (0.46 g, 0.915 mmol) and DMAP (0.056 g, 0.458 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (10 mL). Column chromatography (SiO<sub>2</sub>, cyclohexane/diethyl ether, 9:1) gave II-32a (0.35 g, 61% calc. for two steps) as a colorless oil. IR (neat): ν = 1753 (C=O), 1736 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.25 (d, J = 8.5 Hz, 4 H), 6.87 (d, J = 8.5 Hz, 4 H), 4.43 (s, 4 H), 4.29 (t, J = 6.5 Hz, 4 H), 4.27 (t, J = 6.5 Hz, 4 H), 3.90 (m, 12 H), 3.80 (s, 6 H), 3.58 (t, J = 6 Hz, 4 H), 3.36 (s, 4 H), 1.87 (m, 12 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>): δ = 166.7, 159.3, 130.8, 129.4, 113.9, 72.8, 66.8, 62.8, 62.7, 60.3, 60.2, 55.4, 41.6, 33.2, 31.9, 28.0, 27.9, 21.3 ppm.

**Compound II-32b:** II-32b was synthesized from DCC (0.87 g, 4.20 mmol), II-14b (0.38 g, 1.20 mmol), II-31b and DMAP (0.15 g, 1.20 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (60 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 9:1) gave II-32b (1.70 g, 99%) as a colorless oil. IR (neat): ν = 1753 (C=O), 1736 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): δ = 7.26 (d, J = 8.5 Hz, 4 H), 6.87 (d, J = 8.5 Hz, 4 H), 4.43 (s, 4 H), 4.17 (m, 8 H), 3.82 (m, 18 H), 3.47 (t, J = 6.5 Hz, 4 H), 3.37 (s, 4 H), 1.68 (m, 24 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>): δ = 166.8, 159.3, 130.9, 129.4, 113.9, 72.6, 70.1, 65.6, 65.6, 63.8, 63.3, 63.2, 55.4, 41.7, 29.8, 29.3, 28.0, 27.0, 26.4, 25.3, 21.3 ppm.





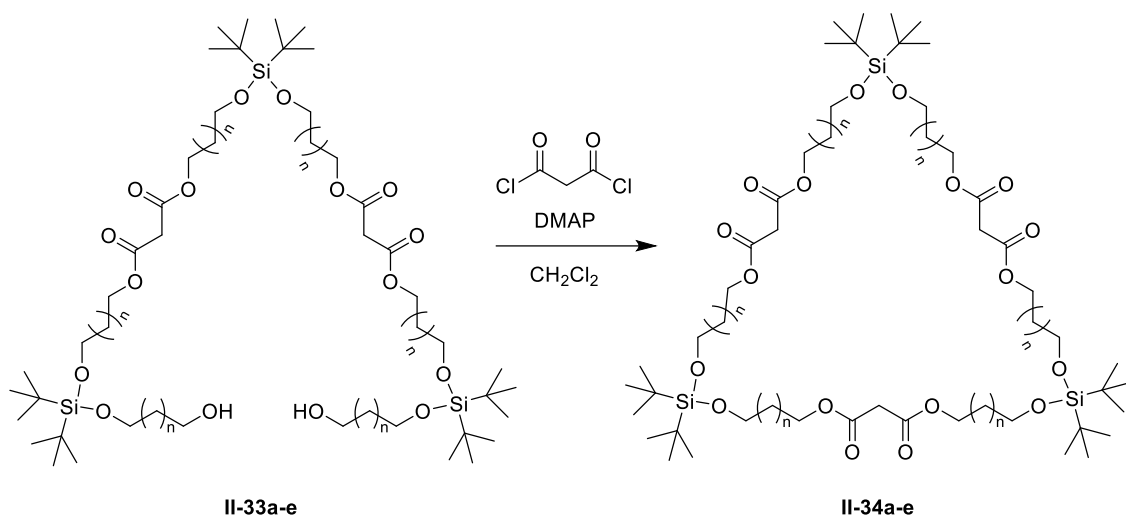
**Compound II-33a:** **II-33a** was synthesized from water (1 mL), **II-32a** (0.348 g, 0.277 mmol) and DDQ (0.139 g, 0.610 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (10 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) and column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 98:2) gave **II-33a** (0.274 g, 98%) as a colorless oil. IR (neat):  $\tilde{\nu}$  = 3459 (O-H), 1752 (C=O), 1737 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.30 (t, *J* = 6.5 Hz, 4 H), 4.29 (t, *J* = 6.5 Hz, 4 H), 4.02 (t, *J* = 5.5 Hz, 4 H), 3.92 (m, 8 H), 3.81 (t, *J* = 6 Hz, 4 H), 3.37 (s, 4 H), 2.05 (s br, 2 H), 1.90 (m, 8 H), 1.81 (m, *J* = 5.5 Hz, 4 H), 1.01 (s, 36 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta$  = 166.8, 166.7, 62.8, 62.8, 62.7, 61.5, 60.4, 60.3, 41.6, 34.8, 31.9, 28.0, 27.9, 21.3 ppm.

**Compound II-33b:** **II-33b** was synthesized from water (1.5 mL), **II-32b** (1.68 g, 1.26 mmol) and DDQ (0.571 g, 2.51 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (80 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 97:3) and column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 97:3) gave **II-33b** (1.23 g, 89%) as a colorless oil. IR (neat):  $\tilde{\nu}$  = 3454 (O-H), 1752 (C=O), 1736 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.19 (t, *J* = 6.5 Hz, 8 H), 3.85 (m, 12 H), 3.67 (t, *J* = 6 Hz, 4 H), 3.38 (s, 4 H), 1.67 (m, 26 H), 1.00 (s, 36 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta$  = 166.8, 65.6, 64.0, 63.3, 62.9, 41.7, 30.0, 29.7, 29.3, 29.2, 28.0, 25.3, 25.2, 21.3 ppm.

**Compound II-33c:** **II-33c** was synthesized from water (5 mL), **II-32c** (2.06 g, 1.45 mmol) and DDQ (0.658 g, 2.90 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (50 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) and column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/ diethyl ether, 8:2) gave **II-33c** (0.916 g, 53%) as a colorless oil. IR (neat):  $\tilde{\nu}$  = 3454 (O-H), 1752 (C=O), 1736 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.15 (t, *J* = 6.5 Hz, 8 H), 3.81 (m, 12 H), 3.65 (t, *J* = 6.5 Hz, 4 H), 3.35 (s, 4 H), 1.58 (m, 36 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta$  = 166.8, 65.7, 63.8, 63.6, 63.1, 41.7, 32.8, 32.7, 32.5, 28.4, 28.0, 22.3, 22.2, 21.3 ppm.

**Compound II-33d:** **II-33d** was synthesized from water (3 mL), **II-32d** (1.26 g, 0.834 mmol) and DDQ (0.417 g, 1.83 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (30 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 99:1) and column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/ diethyl ether, 1:1 and 0.1% MeOH) gave **II-33d** (0.683 g, 65%) as a colorless oil. IR (neat):  $\tilde{\nu}$  = 3418 (O-H), 1752 (C=O), 1737 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.14 (t, *J* = 7 Hz, 8 H), 3.81 (m, 12 H), 3.64 (t, *J* = 6.5 Hz, 4 H), 3.36 (s, 4 H), 1.66 (m, 8 H), 1.56 (m, 16 H), 1.40 (m, 26 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta$  = 166.9, 65.8, 63.9, 63.8, 63.1, 41.8, 33.0, 33.0, 32.9, 28.7, 28.0, 25.8, 25.8, 25.8, 25.7, 25.6, 25.6, 21.3 ppm.

**Compound II-33e:** **II-33e** was synthesized from water (1 mL), **II-32e** (0.744 g, 0.468 mmol) and DDQ (0.213 g, 0.938 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (35 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) and column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 99:1) gave **II-33e** (0.450 g, 71%) as a colorless oil. IR (neat):  $\tilde{\nu}$  = 3454 (O-H), 1753 (C=O), 1737 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.14 (t, *J* = 7 Hz, 8 H), 3.80 (t, *J* = 6.5 Hz, 12 H), 3.64 (t, *J* = 6.5 Hz, 4 H), 3.37 (s, 4 H), 1.65 (m, 8 H), 1.55 (m, 18 H), 1.35 (m, 34 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta$  = 166.9, 65.8, 63.9, 63.9, 63.2, 41.8, 33.0, 33.0, 33.0, 29.4, 29.2, 28.6, 28.0, 26.0, 26.0, 25.9, 21.3 ppm.



**General procedure for the synthesis of compounds II-34a-e:** Malonyl chloride (1.1 eq) was added to a solution of II-33a-e (1 eq) and DMAP (2 eq) in CH<sub>2</sub>Cl<sub>2</sub> under argon. After 24 h the mixture was filtered on SiO<sub>2</sub> and concentrated. Column chromatography (SiO<sub>2</sub>) gave II-34b-e.

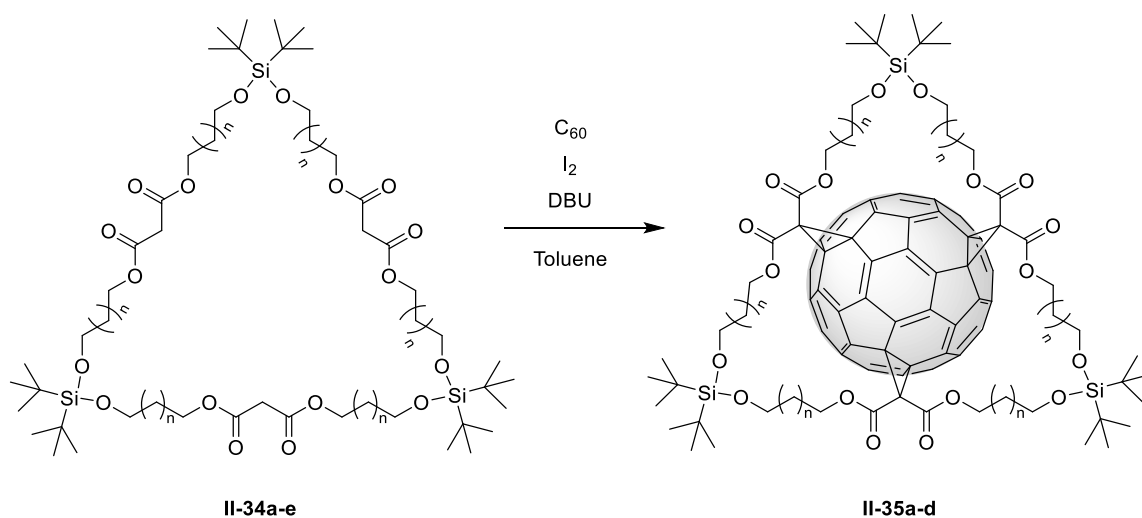
**Compound II-34a:** II-34a was synthesized from malonyl chloride (0.0564 g, 0.400 mmol), II-33a (0.405 g, 0.400 mmol) and DMAP (0.0977 g, 0.800 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (50 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/ diethyl ether, 9:1) and column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/ diethyl ether, 9:1) gave II-34a (0.229 g, 53 %) as a colorless oil. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>): δ = 4.22 (t, *J* = 6 Hz, 12 H), 3.84 (t, *J* = 6 Hz, 12 H), 3.30 (s, 6 H), 1.83 (quint., *J* = 6 Hz, 12 H), 0.93 (s, 54 H) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>): δ = 166.6, 62.5, 60.1, 41.5, 31.7, 27.8, 21.1 ppm. MALDI-TOF-MS: 1103.5 (100%, [M+Na]<sup>+</sup>, calcd for C<sub>51</sub>H<sub>96</sub>O<sub>18</sub>Si<sub>3</sub>Na: 1103.6), 1081.5 (47%, [M+H]<sup>+</sup>, calcd for C<sub>51</sub>H<sub>97</sub>O<sub>18</sub>Si<sub>3</sub>: 1081.6).

**Compound II-34b:** II-34b was synthesized from malonyl chloride (0.156 g, 1.11 mmol), II-33b (1.22 g, 1.11 mmol) and DMAP (0.305 g, 2.50 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (300 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/ EtOAc, 9:1) and column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) gave II-34b (0.331 g, 26 %) as a colorless oil. IR (neat):  $\tilde{\nu}$  = 1752 (C=O), 1735 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): δ = 4.18 (t, *J* = 6.5 Hz, 12 H), 3.84 (t, *J* = 6 Hz, 12 H), 3.37 (s, 6 H), 1.76 (q, *J* = 6.5 Hz, 12 H), 1.61 (t, *J* = 6 Hz, 12 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>): δ = 166.8, 65.5, 63.3, 41.8, 29.3, 28.0, 25.3, 21.3 ppm. MALDI-TOF MS: *m/z* = 1417.740 [M+H]<sup>+</sup>, calcd for C<sub>75</sub>H<sub>145</sub>O<sub>18</sub>Si<sub>3</sub>: 1417.97.

**Compound II-34c:** II-34c was synthesized from malonyl chloride (0.109 g, 0.775 mmol), II-33c (0.916 g, 0.775 mmol) and DMAP (0.189 g, 1.55 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (250 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/diethyl ether, 1:1) and column chromatography (SiO<sub>2</sub>, cyclohexane / diethyl ether, 1:1) gave II-34c (0.612 g, 63%) as a colorless oil. IR (neat):  $\tilde{\nu}$  = 1753 (C=O), 1736 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): δ = 4.15 (t, *J* = 6.5 Hz, 12 H), 3.81 (t, *J* = 6.5 Hz, 12 H), 3.35 (s, 6 H), 1.68 (m, 12 H), 1.58 (m, 12 H), 1.45 (m, 12 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (75 MHz, CDCl<sub>3</sub>): δ = 166.8, 65.7, 63.6, 41.7, 32.5, 28.4, 28.0, 22.8, 21.3 ppm. MALDI-TOF MS: *m/z* = 1271.6 [M+Na]<sup>+</sup>, calcd for C<sub>63</sub>H<sub>120</sub>NaO<sub>18</sub>Si<sub>3</sub>: 1271.77.

**Compound II-34d:** **II-34d** was synthesized from malonyl chloride (0.0835 g, 0.592 mmol), **II-33d** (0.682 g, 0.539 mmol) and DMAP (0.132 g, 1.08 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (70 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/diethyl ether, 1:1) and column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/ diethyl ether, 7:3) gave **II-34d** (0.307 g, 43%) as a colorless oil. IR (neat):  $\tilde{\nu}$  = 1753 (C=O), 1736 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.14 (t, *J* = 6.5 Hz, 12 H), 3.80 (t, *J* = 6 Hz, 12 H), 3.36 (s, 6 H), 1.66 (m, 12 H), 1.55 (m, 12 H), 1.39 (m, 24 H), 0.99 (s, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta$  = 166.8, 65.7, 63.8, 41.8, 32.9, 28.7, 28.0, 25.8, 25.6, 21.3 ppm. MALDI-TOF MS: *m/z* = 1355.8 [M+Na]<sup>+</sup> calcd for C<sub>69</sub>H<sub>132</sub>NaO<sub>18</sub>Si<sub>3</sub>: 1355.86.

**Compound II-34e:** **II-34e** was synthesized from malonyl chloride (0.0470 g, 0.333 mmol), **II-33e** (0.450 g, 0.333 mmol) and DMAP (0.122 g, 1.00 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (200 mL). Filtration (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) and column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) gave **II-34e** (0.099 g, 21%) as a colorless oil. <sup>1</sup>H NMR (300 MHz, CDCl<sub>3</sub>):  $\delta$  = 4.17 (t, *J* = 6.5 Hz, 12 H), 3.83 (t, *J* = 6.5 Hz, 12 H), 3.40 (s, 6 H), 1.62 (m, 24 H), 1.39 (m, 36 H), 1.03 (m, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta$  = 166.7, 65.7, 63.8, 41.8, 32.9, 29.1, 28.5, 28.0, 25.9, 25.8, 21.2 ppm. MALDI-TOF MS: *m/z* = 1417.740 [M+H]<sup>+</sup>, calcd for C<sub>75</sub>H<sub>145</sub>O<sub>18</sub>Si<sub>3</sub>: 1417.97.



**General procedure for the synthesis of compounds II-35a-d:** DBU (7.5 eq) was added to a solution of **II-34** (1 eq), [60]fullerene (C<sub>60</sub>) (1 eq) and I<sub>2</sub> (3.5 eq) in toluene. The mixture was stirred at room temperature for 1 h 30 min, then filtered on SiO<sub>2</sub> (cyclohexane, then CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography (SiO<sub>2</sub>) followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) gave **II-35a-d**.

**Compound II-35a:** **II-35a** was synthesized from DBU (0.15 mL, 1.00 mmol), **II-34a** (0.58 g, 0.15 mmol), C<sub>60</sub> (0.108 g, 0.15 mmol) and I<sub>2</sub> (0.131 g, 0.52 mmol) in toluene (400 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 1:1) gave **II-35a** as a cherry-red glassy solid (tris-adduct *e,e,e*, 0.055 g, 61%). IR (neat):  $\tilde{\nu}$  = 1751 (C=O) cm<sup>-1</sup>; UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>): 252 (1.6x10<sup>5</sup>), 281 (1.1x10<sup>5</sup>), 304 (sh, 9x10<sup>4</sup>), 482 (6x10<sup>4</sup>), 564 (sh, 2x10<sup>4</sup>); <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  = 4.70 (m, 3 H), 4.60 (m, 3 H), 4.36 (m, 6 H), 3.95-3.77 (m, 12 H), 1.94 (m, 12 H), 0.98 (s, 36 H), 0.90 (s, 36

H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz): δ = ppm. MALDI-TOF-MS: 1797.0 ([M+H]<sup>+</sup>, calcd for C<sub>98</sub>H<sub>91</sub>O<sub>18</sub>Si<sub>3</sub>: 1796.5).

**Compound II-35b:** **II-35b** was synthesized from DBU (0.299 mL, 1.96 mmol), **II-34b** (0.326 g, 0.280 mmol), C<sub>60</sub> (0.201 g, 0.280 mmol) and I<sub>2</sub> (0.248 g, 0.978 mmol) in toluene (400 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 1:1) gave **II-35b** as a cherry-red glassy solid (tris-adduct *e,e,e*, 0.204 g, 39%). IR (neat):  $\tilde{\nu}$  = 1748 (C=O), 1728 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>): 237 (1.2x10<sup>5</sup>), 251 (1.2x10<sup>5</sup>), 281 (9x10<sup>4</sup>), 304 (sh, 7x10<sup>4</sup>), 484 (6x10<sup>3</sup>), 568 (sh, 2x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): δ = 4.53 (m, 6 H), 4.21 (m, 6 H), 3.79 (m, 12 H), 1.77 (m, 12 H), 1.57 (m, 12 H), 1.00 (s, 27 H), 0.96 (s, 27 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz): δ = 163.6, 163.2, 147.8, 146.8, 146.8 (two peaks), 146.8, 146.6, 146.6, 146.5, 145.8, 145.7, 144.7, 144.6, 144.2, 143.7, 142.7, 142.5, 142.4, 141.7, 141.2, 71.0, 70.4, 67.3, 67.1, 63.2, 62.9, 53.9, 29.5, 29.2, 28.0, 25.5, 25.4, 21.4, 21.2 ppm. MALDI-TOF-MS: 1881.7 ([M+2H]<sup>+</sup>, calcd for C<sub>117</sub>H<sub>104</sub>O<sub>18</sub>Si<sub>3</sub>: 1881.3).

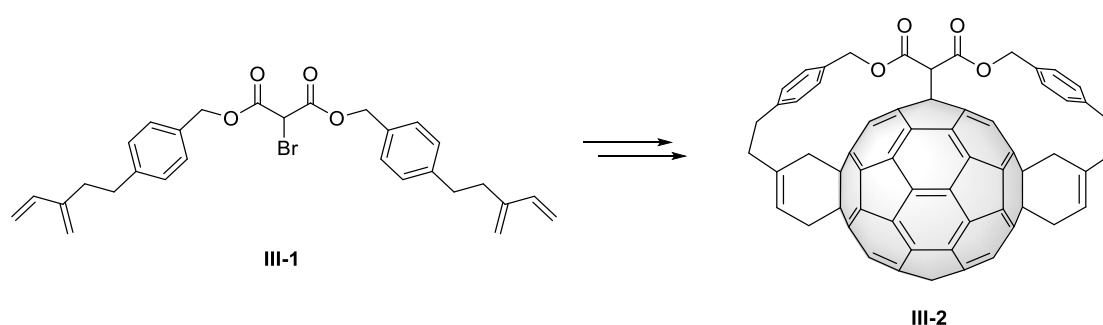
**Compound II-35c:** **II-35c** was synthesized from DBU (0.557 mL, 3.68 mmol), **II-34c** (0.326 g, 0.280 mmol), C<sub>60</sub> (0.353 g, 0.490 mmol) and I<sub>2</sub> (0.435 g, 1.72 mmol) in toluene (700 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 1:1) followed by gel permeation chromatography gave **II-35c** as a cherry-red glassy solid (tris-adduct *e,e,e*, 77 mg, 8%). IR (neat):  $\tilde{\nu}$  = 1746 (C=O) cm<sup>-1</sup>. UV/vis (CH<sub>2</sub>Cl<sub>2</sub>): 239 (9x10<sup>4</sup>), 250 (9x10<sup>4</sup>), 281 (6x10<sup>4</sup>), 307 (sh, 5x10<sup>4</sup>), 484 (4x10<sup>3</sup>), 564 (sh, 1x10<sup>3</sup>) nm. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>): δ = 4.49 (m, 2 H), 4.19 (m, 2 H), 3.81-3.70 (m, 4 H), 1.70 (m, 4 H), 1.55 (m, 4 H), 1.41 (m, 4 H), 0.98 (s, 18 H) ppm. <sup>13</sup>C NMR (75 MHz, CDCl<sub>3</sub>): δ = 163.6, 163.6, 147.8, 146.9, 146.8, 146.8, 146.6, 146.5, 146.5, 145.8, 145.8, 144.8, 144.6, 144.5, 143.7, 143.0, 142.4, 142.3, 141.8, 141.0, 71.1, 70.4, 67.3, 67.2, 63.5, 63.4, 53.7, 32.3, 32.2, 28.3, 28.1, 28.0, 22.2, 22.1, 21.3, 21.2 ppm. MALDI-TOF-MS: 1963.942 ([M+H]<sup>+</sup>, calcd for C<sub>123</sub>H<sub>114</sub>O<sub>18</sub>Si<sub>3</sub>: 1962.73).

**Compound II-35d:** **II-35d** was synthesized from DBU (0.250 mL, 1.65 mmol), **II-34d** (0.293 g, 0.219 mmol), C<sub>60</sub> (0.158 g, 0.219 mmol) and I<sub>2</sub> (0.195 g, 0.768 mmol) in toluene (316 mL). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 1:1) followed by gel permeation chromatography gave **II-35d** as a cherry-red glassy solid (tris-adduct *e,e,e*, 25.4 mg, 5.7%). IR (neat):  $\tilde{\nu}$  = 1748 (C=O) cm<sup>-1</sup>. UV/vis (CH<sub>2</sub>Cl<sub>2</sub>): 242 (1.0x10<sup>5</sup>), 248 (1.0x10<sup>5</sup>), 285 (sh, 6x10<sup>4</sup>), 483 (4x10<sup>3</sup>), 564 (sh, 1x10<sup>3</sup>) nm. <sup>1</sup>H NMR (300 MHz, CDCl<sub>3</sub>): δ = 4.70 (m, 4 H), 4.41 (m, 4 H), 4.23 (m, 4 H), 3.78 (m, 12 H), 1.70 (m, 12 H), 1.39 (m, 36 H), 0.99 (m, 54 H) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>): δ = 163.6, 163.5, 147.6, 146.9, 146.9, 146.7, 146.6, 146.6, 146.5, 145.9, 145.8, 144.8, 144.5 (two peaks), 143.6, 143.0, 142.5, 142.3, 141.9, 141.0, 129.2, 128.4, 125.4, 71.1, 70.4, 67.1, 67.1, 63.7, 63.7, 53.6, 32.7, 28.6, 28.0, 25.6, 25.5, 25.4, 25.2, 21.3, 21.3 ppm. MALDI-TOF MS: *m/z* = 2047.948 [M+H]<sup>+</sup>, calcd for C<sub>129</sub>H<sub>127</sub>O<sub>18</sub>Si<sub>3</sub>: 2047.83

# **Chapter III: Regioselective syntheses of C<sub>60</sub> Tris-adducts by a tethered-directed approach.**

## 1. Introduction.

The first regioselective formation of a  $C_{60}$  tris-adduct was reported in 1994 by Diederich and coworkers, introducing the tethered-directed remote functionalization concept.<sup>[1]</sup> Their methodology was based on three elements: (1) an anchor; (2) a tether and (3) a reactive group. The anchor is the first addend attached to the  $C_{60}$  core. It is linked to the reactive groups by the tethers. The length and the shape of the tether will then induce the regioselective attack of the reactive group onto the  $C_{60}$  core to form regioselectively a  $C_{60}$  derivative. A malonate moiety was chosen as anchor, a 4-substituted benzyl alcohol as tether and a 2-substituted 1,3-butadiene as reactive group. Reaction of **III-1** with  $C_{60}$  in a two steps synthesis led only to the tris-adducts *e,e,trans-1* **III-2** in an excellent yield of 48% (Figure III-1).



**Figure III-1** : First regioselective synthesis of  $C_{60}$  tris-adducts by tethered-directed remote functionalization.

This tether-directed approach has been also extensively used for the regioselective synthesis of  $C_{60}$  bis-adducts.<sup>[2]</sup> Based on double Bingel reactions, 7 out of the 8 possible regioisomers of  $C_{60}$  bis-adducts could be obtained. These syntheses were achieved with xyllylene,<sup>[3]</sup> crown ether,<sup>[4]</sup> porphyrin,<sup>[5]</sup> Tröger's base,<sup>[6]</sup> phenantroline<sup>[3]</sup>, spirobifluorene<sup>[3]</sup> and chiral<sup>[7]</sup> tethers (Figure III-2). For steric reasons, this strategy is not suitable for the preparation of the *cis-1* regioisomer.

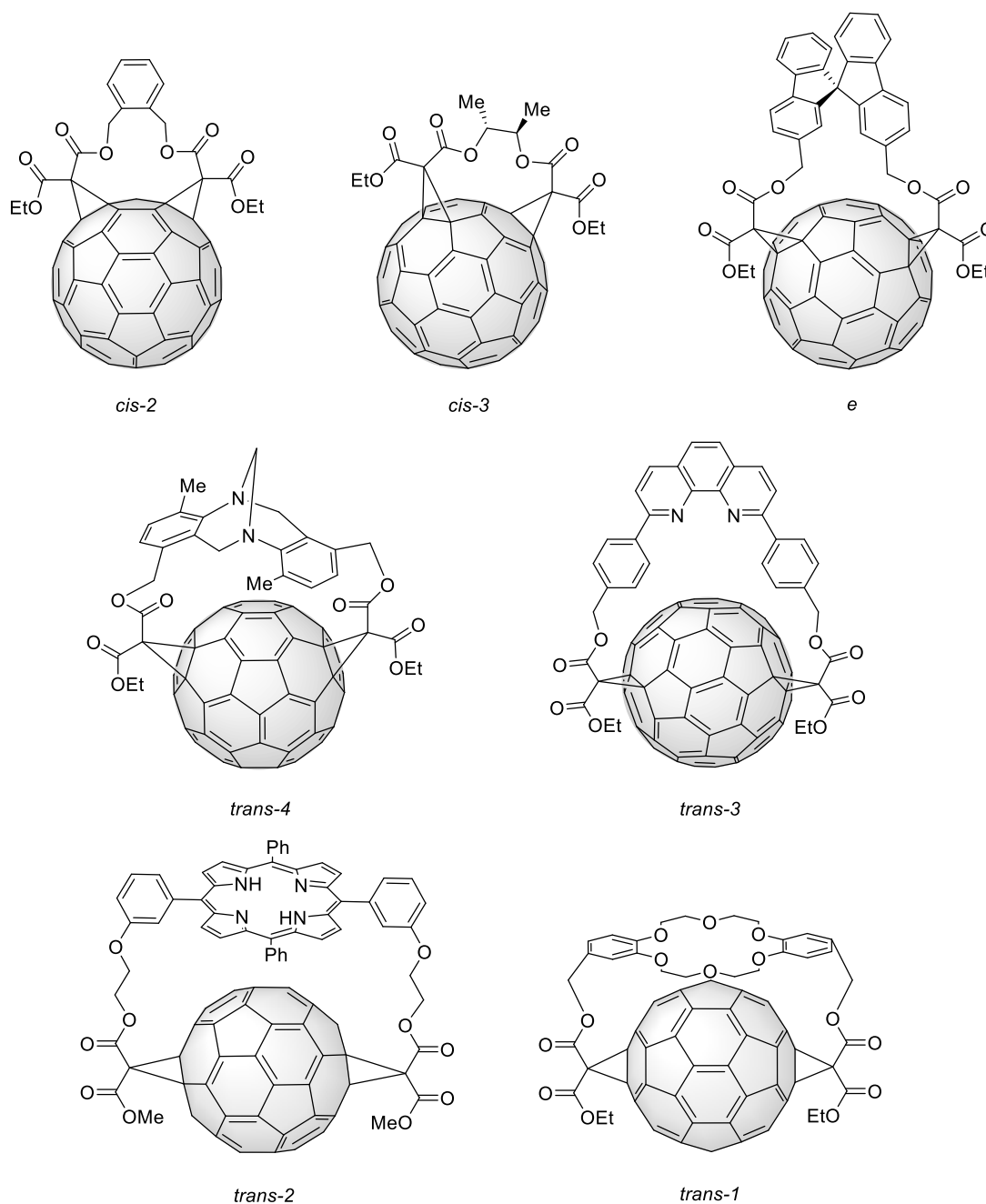


Figure III-2 : Examples of tethers for the regioselective bis-functionalization of  $C_{60}$ .

Theoretically, it is more complicated to obtain  $C_{60}$  tris-adducts in a pure form due to the 46 possible addition patterns. This problem has been partially solved by the tether-directed remote functionalization methodology as the number of possible regioisomers is considerably reduced. Different tethers have been developed in order to prepare selected fullerene tris-adducts.

Diederich and coworkers reported in 1999 a tether-directed methodology based on a cyclotrimeratrylene (CTV) tether.<sup>[8]</sup> The CTV core was selected for its favorable  $\pi$ - $\pi$  charge transfer interaction with  $C_{60}$  and the resulting host-guest complex CTV- $C_{60}$  may play the role of a template.<sup>[9]</sup> Subsequent  $C_{60}$  regioselective functionalization was enabled by grafting



malonates on the CTV and afforded two pairs of diastereomeric *trans-3,trans-3,trans-3* tris-adducts in 11% and 9% yields, respectively (Figure III-3).<sup>[8,10]</sup> Due to the chirality of the CTV tether, the unambiguous assignment of the compounds was difficult. In opposition to the previously reported tris-adducts,<sup>[11]</sup> the formation of those tris-adducts were achieved in a one-step synthesis and isolated by classical silica column chromatography.

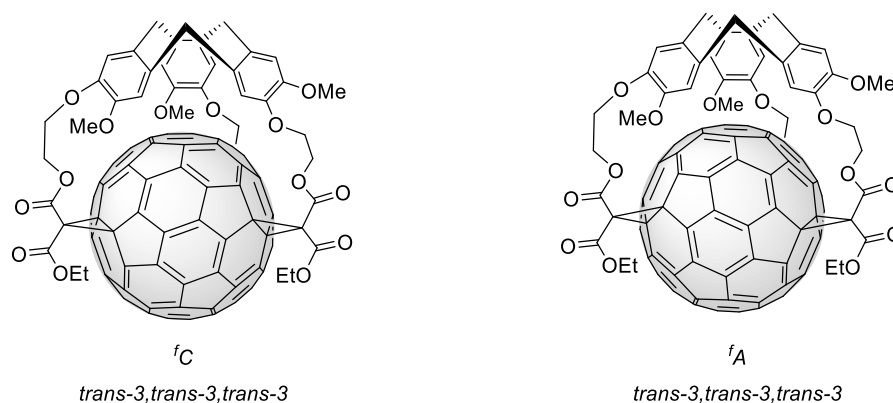


Figure III-3 : Regioselective tris-functionalization of fullerene with a CTV tether. Only one pair of diastereoisomers is showed.

Hirsch and coworkers reported a regioselective synthesis of *e,e,e* tris-adducts with a tripodal tether that could be removed after the formation of the tris-adducts.<sup>[12,13]</sup> In a 7-steps synthesis, they obtained a tripodal benzyl tether bearings three malonates. Subsequent reaction of the tripodal malonates with C<sub>60</sub> afforded the *e,e,e* tris-adducts in addition to an inseparable mixture of other tris-adducts. Deprotection of the benzyl tether was then achieved to obtain the resulting tris-adducts triol or tribromide that allowed further functionalization of the tris-adducts (Figure III-4). Furthermore, the regioselectivity could be increased by using another tripodal tether with shorter distances between the tether and the malonates moieties. This result showed the influence of the spacer between the tether and the malonates moieties on the regioselectivity. Moreover, they also modified the side chain of the malonate moieties in order to obtain tris-adducts with functionalizable side chains. In addition to being regioselective, this methodology is also stereoselective. Indeed, it provides two defined addend zones. However, the high number of steps for the preparation of the tripodal tether discouraged the use of this methodology.

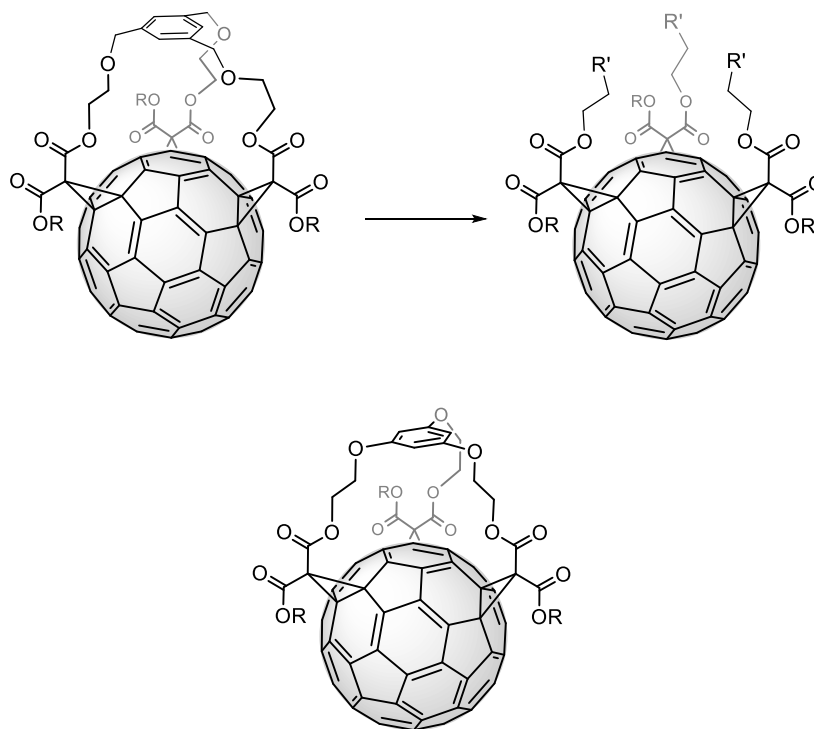


Figure III-4: (top) Deprotectable tether allowing further functionalization. (bottom) Non-deprotectable tether which afford better regioselectivity.

Chronakis and coworkers designed the structurally new tris-malonates tether **III-3** in which two malonates are included in two crown ether type macrocycles. Reaction of **III-3** with  $C_{60}$  gave only the *e,e,trans-1* tris-adducts **III-4** in an excellent yield of 65% (Figure III-5). The assignation of the addition pattern was supported by an X-Ray crystal structures.

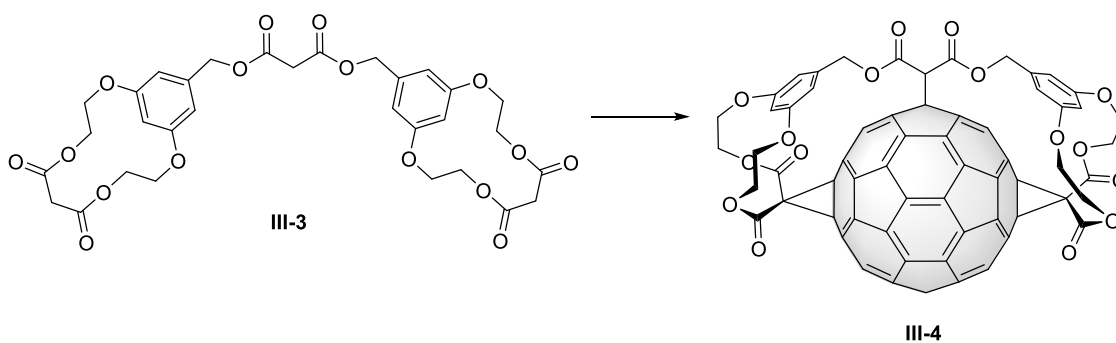


Figure III-5: Original tether for the regioselective functionalization of  $C_{60}$ .

Rubin and Qian proposed a different approach for the regioselective formation of *e,e,trans-1* tris-adducts.<sup>[14]</sup> They designed a removable tether that led to the formation of *trans-1* bis-adducts via a double Diels-Alder reaction. This  $C_{60}$  bis-adduct served as a template. Indeed, this *trans-1* addition pattern has a high directing effects for the *equatorial* positions.<sup>[15]</sup> By masking an *equatorial* position with the tether, the functionalization of the three remaining

equatorial positions afforded a temporary penta-adducts **III-5** (Figure III-6). The tris-adduct **III-6** was obtained by removal of the tether by a retro-Diels-Alder procedure.

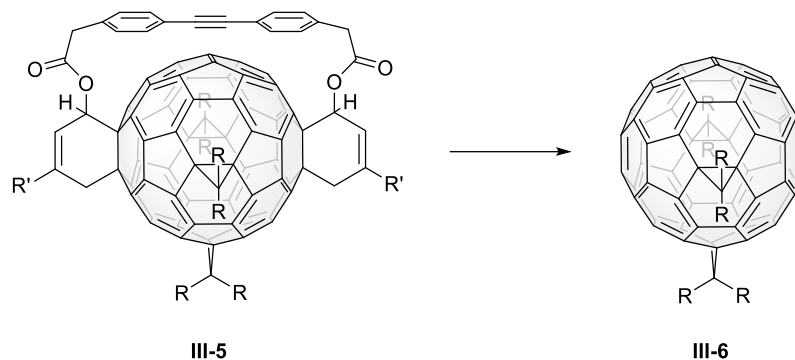


Figure III-6 : Formation of tris-adducts with a removable tether as double bonds activator.

With a similar approach, Rubin's group have synthesized regioselectively a *trans-4,trans-4,trans-4* tris-adduct.<sup>[16]</sup> First, they synthesized a *cis-1*  $C_{60}$  bis-adducts with a removable tether. This intermediate had three activated double bonds in a *trans-4* positional relationships. Secondly, functionalization of these activated double bonds and subsequent removal of the tether afforded the *all-trans-4* tris-adducts (Figure III-7).

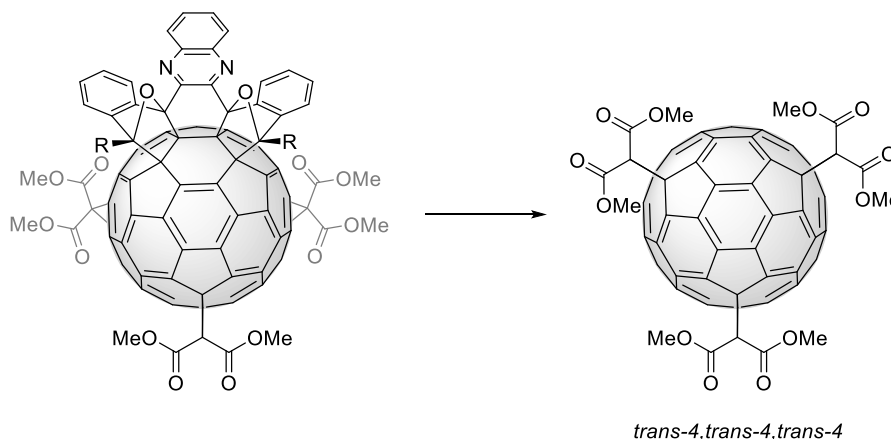


Figure III-7 : Formation of the *all-trans-4* tris-adducts by temporary saturation of the *cis-1* positions.

In the reported methodologies, either the starting tris-malonates precursors are rather difficult to synthesize or the bridging subunits allowing the control of the addition pattern are difficult to remove upon reaction with the fullerene, thus limiting the accessibility of fullerene tris-adducts building blocks. In this context, our group became interested in the development of a quite simple tether-directed methodology for the formation of  $C_{60}$  *e,e,e* tris-adducts. In this effect, the two important criteria were an efficient synthesis with a low number of steps for the formation of the tris-malonates tether and an easily deprotectable tether. Based on a silyl tether linked to the malonates by ethyl spacers, the *e,e,e* tris-adducts was regioselectively obtained in three steps from commercial starting materials.<sup>[17]</sup> The *e,e,e* addition pattern was

confirmed by spectroscopic data and by a X-Ray crystal structure. The silyl tether was easily removed to afford the corresponding tris-adducts triol.

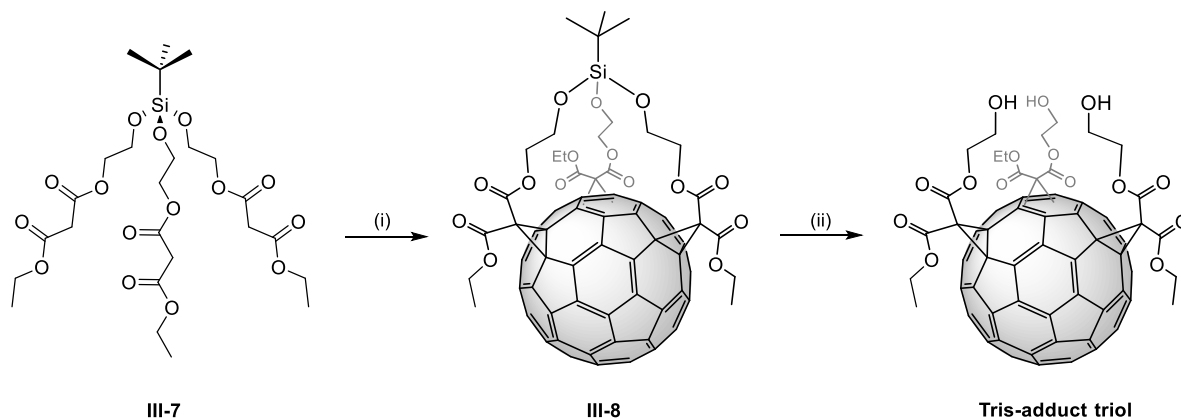


Figure III-8 : Preliminary results for the regioselective functionalization of  $C_{60}$  based on a trialkoxysilane tether.

However, the 8% yield for the threefold Bingel reaction of the tris-malonates tether with  $C_{60}$  is not satisfying and prevents a large scale synthesis. In this chapter, the preparation of *e,e,e* tris-adducts with different spacers between the silyl tether and the malonates will be presented. Our aim being to optimize the yield for the preparation of fullerene *e,e,e* tris-adducts.

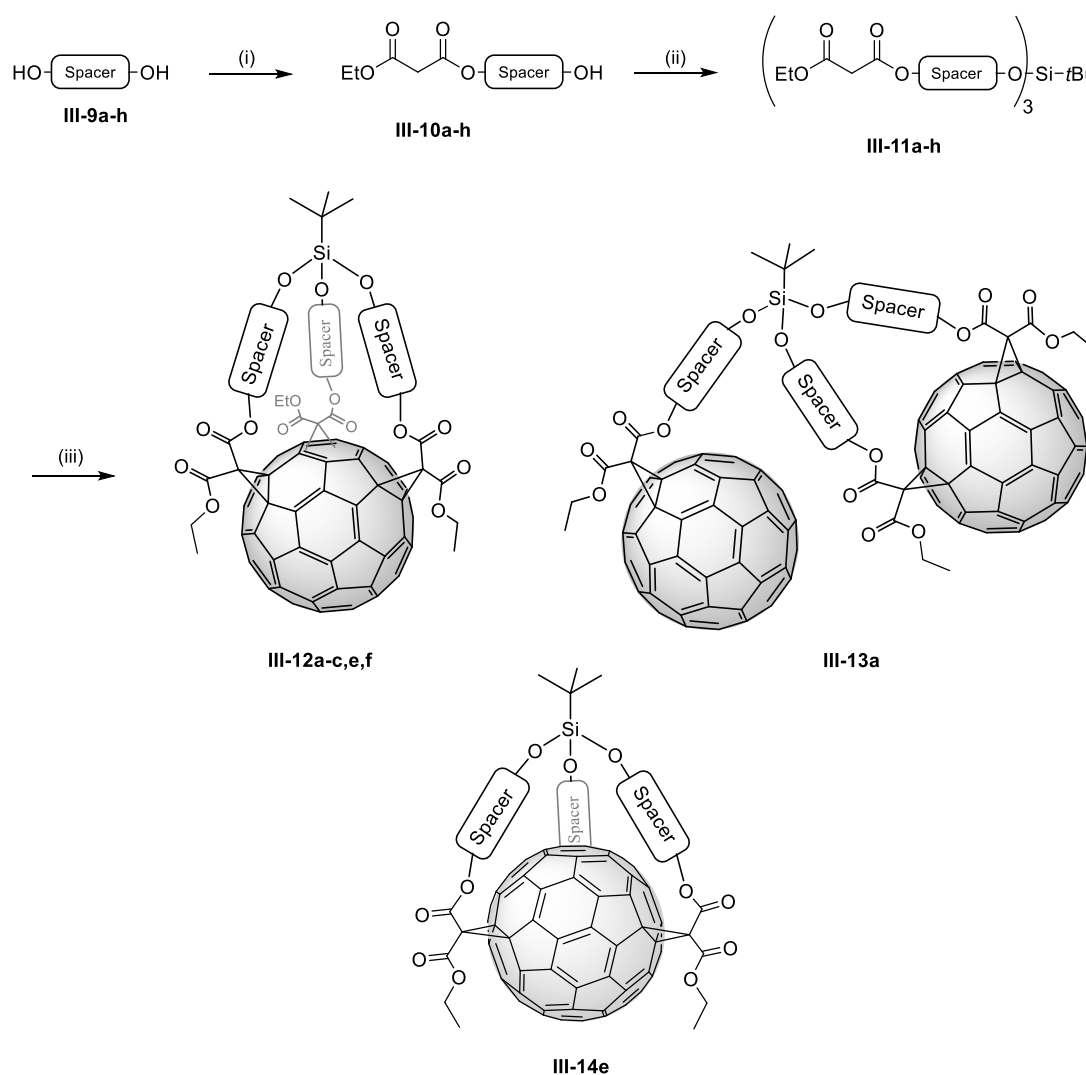
## 2. Regioselective syntheses of $C_{60}$ Tris-adducts.

### 2.1 Syntheses.

As shown in the literature, the tether/spacer that link the malonates determine the regioselectivity. The silane tether developed in the group has shown promising results for the regioselective functionalization of  $C_{60}$ . In order to increase the yield of the formation of the  $C_{60}$  *e,e,e* tris-adducts, 8 different spacers were tested (Table III-1).

The preparation of the tris-malonate precursors and their subsequent reactions with  $C_{60}$  are depicted in Scheme III-1. Esterification of ethyl malonyl chloride with diols III-9a-h gave malonates III-10a-h. By using a large excess of diol, the formation of the corresponding bis-malonates are largely prevented and compounds III-10a-h were thus obtained in moderate to good yields. Treatment of III-10a-h with tert-butyl(trichloro)silane ( $t\text{-BuSiCl}_3$ ) in dry DMF in the presence of imidazole gave the tris-malonates III-11a-h. Compounds III-11a-h were found to be only moderately stable but their purifications by column chromatography on  $\text{SiO}_2$  were possible. Rapid decomposition was however observed upon storage even at low temperature. Compounds III-11a-h should be used for the next step within the next 12 h following their purifications.

The reactions of **III-11** with  $C_{60}$  were performed under the conditions developed in our laboratory for the preparation of fullerene *e,e,e* tris-adducts **III-8** from *t*-butyl(trialkoxo)silane derivative **III-7**. Specifically, DBU was added to a solution of **III-11**,  $C_{60}$  and iodine in toluene at  $-15^{\circ}\text{C}$ . A key feature of these conditions is a strict control of the temperature. Indeed, fast decomposition of the *t*-butyl(trialkoxo)silane precursor was observed when the reaction was performed at a temperature higher than  $0^{\circ}\text{C}$ . Results of the threefold Bingel reactions of compound **III-11a-h** with  $C_{60}$  are shown in Table III-1. In all the cases except for **III-11e**, the fullerene tris-adduct with an *e,e,e* addition pattern was the only isolable regioisomer. The *e,e,e* addition pattern being chiral, the tris-adducts **III-12** were obtained as a racemic.



**Scheme III-1.** Reagents and conditions: (i) Ethyl malonyl chloride, pyridine, THF, rt, 2 h; (ii) *t*BuSiCl<sub>3</sub>, imidazole, DMF, rt, 12 h; (iii)  $C_{60}$ , I<sub>2</sub>, DBU, toluene,  $-15^{\circ}\text{C}$ , 1 h.

In the case of **III-11a**, the reaction was highly regioselective and no regioisomeric tris-adduct of **III-12a** could be obtained. The by-product of the reaction, compound **III-13a** was identified by MALDI-TOF mass spectrometry as a bis-fullerene derivative resulting from the reaction of **III-11a** with two molecules of  $C_{60}$ . The formation of the bis-fullerene **III-13** resulted

from the possibility of the tris-malonate **III-11** to functionalize several molecules of C<sub>60</sub> but also from a stereochemical problem attributable to the tether. In fact, after the second malonate addition, chirality is introduced by the formation of a macrocycle on the C<sub>60</sub> core and thus a stereogenic center is generated on the silicium atom. Four stereoisomers are possible based on the generated chirality (Figure III-9). In the upper cases, the third malonate is directed toward the C<sub>60</sub> core and the intramolecular reactions furnished racemic *e,e,e* tris-adducts. In the lower cases, the third malonate is directed toward the outside. The intramolecular reaction is no longer possible and the intermolecular reaction afforded the bis-fullerene derivatives.

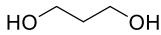
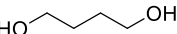
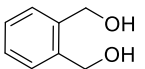
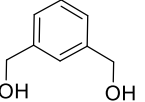
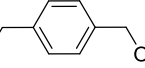
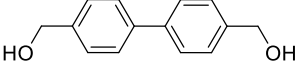
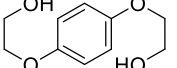

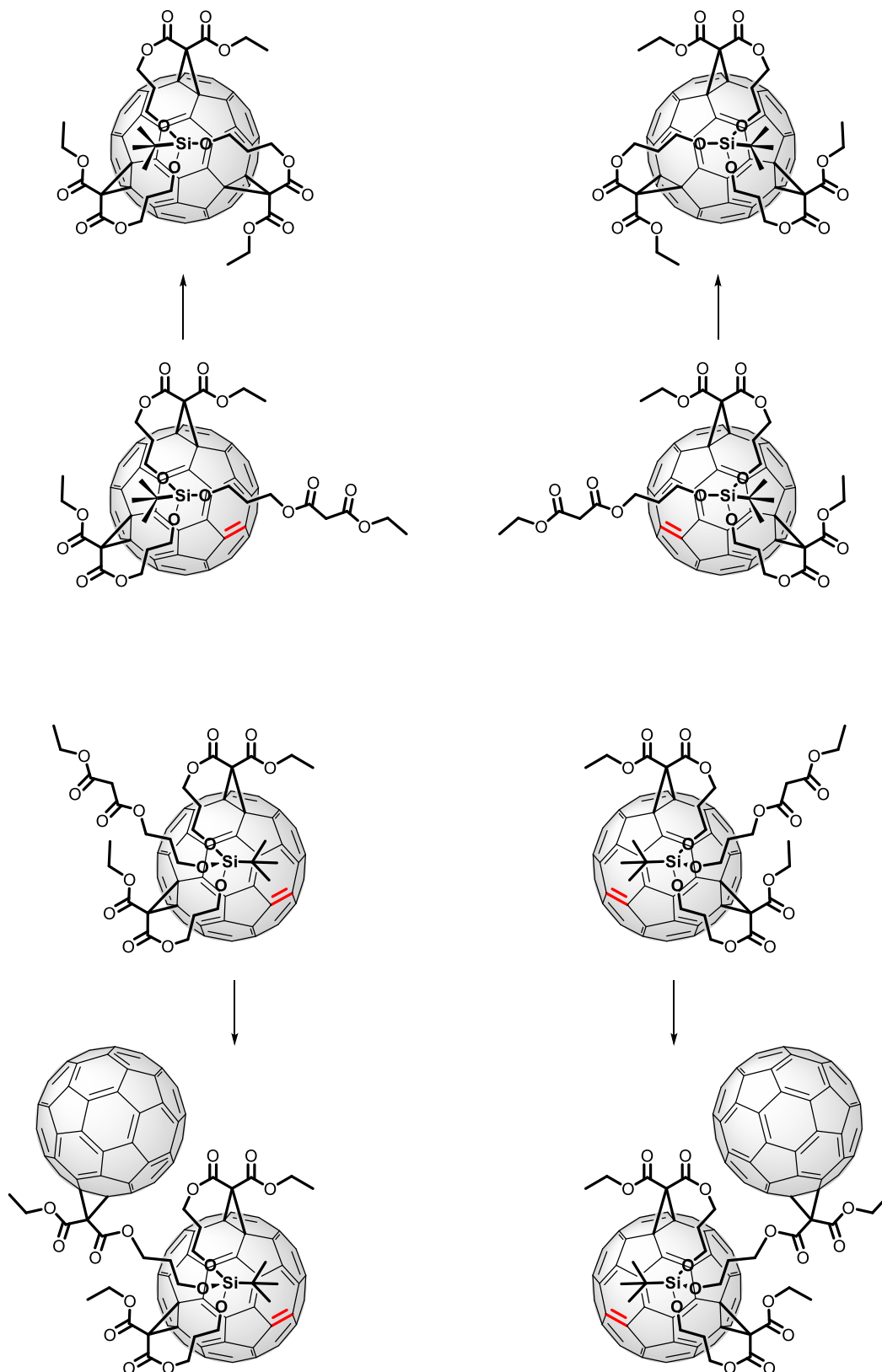
Spacer	Tris-adduit (%)
<b>III-9a</b> 	<b>III-12a</b> 26% <i>e,e,e</i> <b>III-13a</b> 11%
<b>III-9b</b> 	<b>III-12b</b> 14% <i>e,e,e</i>
<b>III-9c</b> 	<b>III-12c</b> 23% <i>e,e,e</i>
<b>III-9d</b> 	<b>Mixture</b>
<b>III-9e</b> 	<b>III-12e</b> 10% <i>e,e,e</i> <b>III-14e</b> 1% <i>all-trans-3</i>
<b>III-9f</b> 	<b>III-12f</b> 3% <i>e,e,e</i>
<b>III-9g</b> 	<b>Mixture</b>
<b>III-9h</b> 	<b>Degradation</b>

Table III-1: Results of the threefold Bingel reactions of the tris-malonates **III-11a-h** with C<sub>60</sub>.

The bis-fullerenes derivatives **III-13** were also formed with the other tris-malonates derivatives **III-11b-g** but were not isolated. For **III-11d** and **III-11g**, no regioselectivity was observed and inseparable mixtures of regioisomers were obtained. In the case of **III-11h**, the silyl tether was too unstable even at -15°C and only degradation products were obtained.

The reactions using compounds with aromatic spacers (**III-11c-e**) showed that the *ortho*, *meta* and *para* orientations have an impact on the outcome of the threefold Bingel reactions with C<sub>60</sub>. The *ortho* derivative **III-11c** furnished the better yield of C<sub>60</sub> *e,e,e* tris-

adducts with a rigid spacer. The meta derivative **III-11d** showed no regioselectivity. On the other hand, the para compound **III-11e** is the only tris-malonate derivative where another regioisomer of C<sub>60</sub> tris-adduct (**III-14e**) was also isolated.

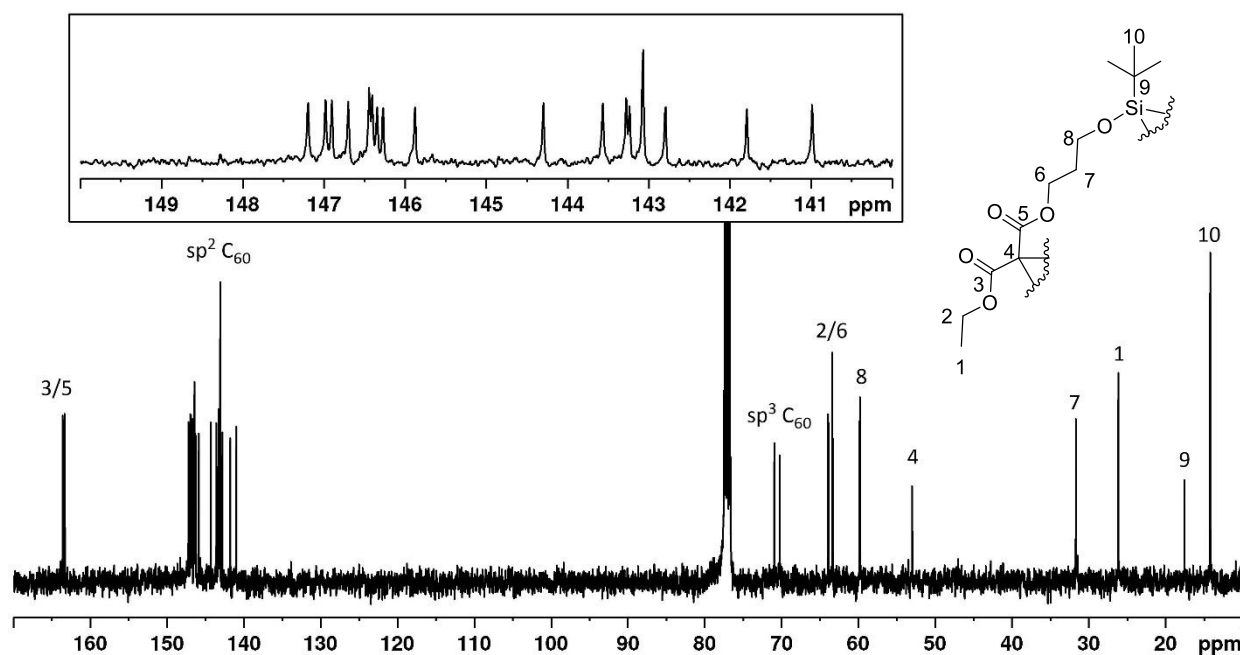


*Figure III-9 : Illustration of the four possible stereoisomers of the intermediary bis-adducts and theirs resulting products.*

All these experiments show that the spacer between the malonate subunits and the central t-butylsilane group plays an important role in the outcome of the reaction as it acts as a directing tether. 1,3-Propane is the most appropriate spacer. Indeed, the best yields in fullerene tris-adducts were obtained when the reaction was performed from **III-11a**.

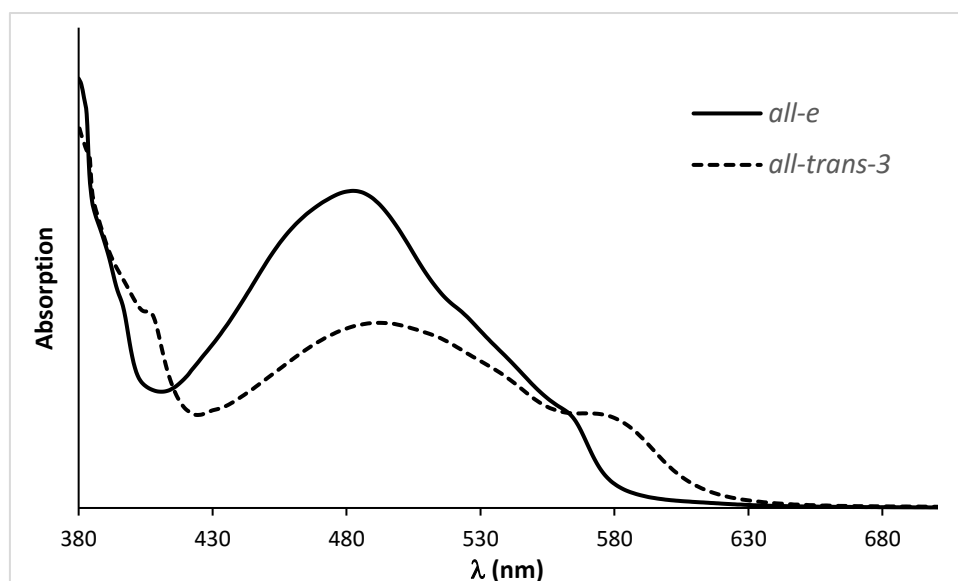
Compounds **III-12a-c,e,f**, **III-13a** and **III-14d** have been characterized by  $^1\text{H}$  and  $^{13}\text{C}$  NMR, UV-vis and IR spectroscopies and MALDI-TOF mass spectrometry. The relative position of the three cyclopropane rings on the  $C_{60}$  core in **III-12** has been determined based on the  $C_3$  molecular symmetry deduced from the  $^1\text{H}$  and  $^{13}\text{C}$  NMR spectra. As a typical example, the  $^{13}\text{C}$  NMR spectrum of **III-12a** recorded in  $\text{CDCl}_3$  is shown in [Figure III-10](#). The expected fullerene resonances are clearly observed: two at  $\delta = 70.2$  and  $70.9$  ppm for the two different  $\text{sp}^3$  C atoms, 17 between  $\delta = 140$  and  $148$  ppm for the different  $\text{sp}^2$  C atoms. All the resonances seen in the  $\text{sp}^2$  region show the same intensity except one signal corresponding to two C atoms having fortuitously the same chemical shift. This is in perfect agreement with the  $C_3$  symmetry of an *e,e,e* fullerene tris-adducts. This is further supported by the 10 expected non-fullerene signals seen for **III-12a**. The cherry-red color and the UV/vis spectrum of **III-12** ([Figure III-11](#)) are also fully consistent with those of previously prepared compound **III-8**.

For **III-13a**, no addition pattern of the bis-adduct part could be assigned based on the  $^1\text{H}$  and  $^{13}\text{C}$  NMR spectra and the UV/vis absorption spectrum.



[Figure III-10](#) :  $^{13}\text{C}$  NMR spectrum (100 MHz,  $\text{CDCl}_3$ ) of **III-12** showing its threefold symmetry. Inset: detailed view showing the resonances of the fullerene  $\text{sp}^2$  C atoms.

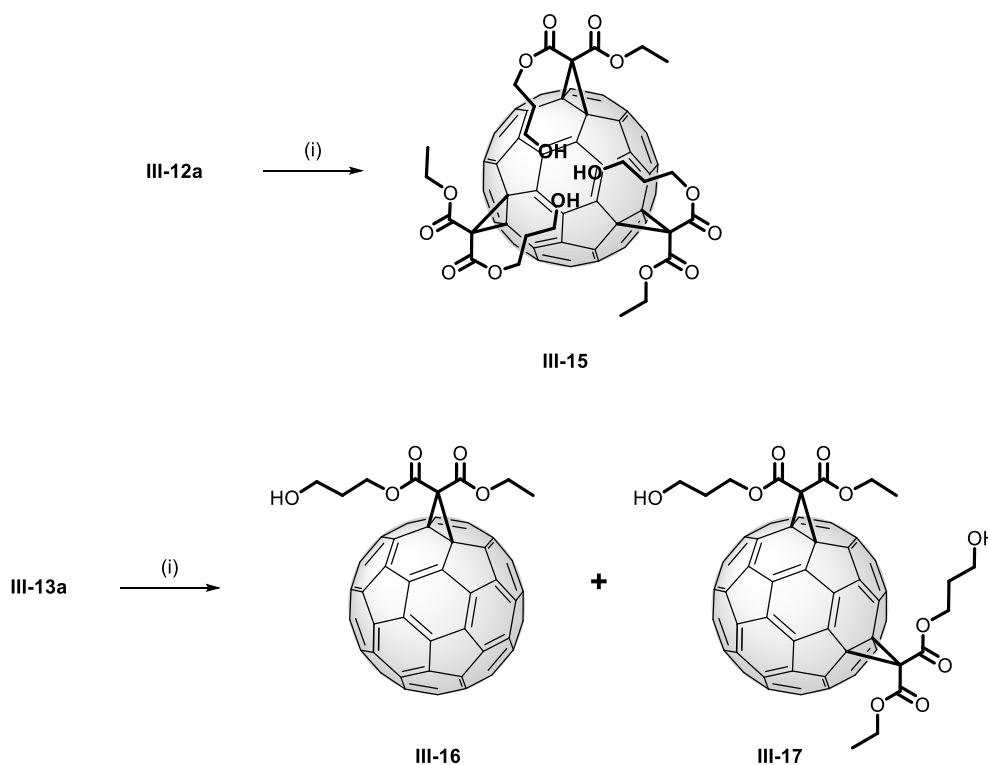




**Figure III-11 :** UV/vis spectrum of **III-12** (—) characteristic of the *e,e,e* addition pattern and UV/vis spectrum of **III-14d** (- - -) characteristic of the *all-trans-3* addition pattern .

The *all-trans-3* tris-adduct **III-14d** is also obtained as a racemic. The *trans-3,trans-3,trans-3* addition pattern was determined based on the  $C_3$  molecular symmetry deduced from the  $^1\text{H}$  and  $^{13}\text{C}$  NMR spectra. It's to note that the  $D_3$  symmetry of the *all-trans-3* addition pattern is reduced by the tether and thus the *all-trans-3* tris-adducts is the only other possible  $C_3$ -symmetrical regioisomer. The UV/vis absorption spectrum, being highly dependent on the addition pattern, was also recorded (**Figure III-11**). Indeed the UV/vis spectrum of **III-14d** was fully consistent with those previously reported for analogous *all-trans-3*  $C_{60}$  tris-adducts.<sup>[18]</sup>

Finally, the removal of the silyl tether has been performed. Basic deprotection conditions are excluded to preserve the ester functions. Our first attempts were then with acidic conditions. The use of trifluoroacetic acid (TFA) or HBr (aq) were inefficient for the deprotection of the silyl tether. It was finally found that the treatment of the silylated fullerene derivatives with an excess of  $\text{BF}_3 \cdot \text{Et}_2\text{O}$  (20 equiv) in  $\text{CH}_2\text{Cl}_2/\text{CH}_3\text{CN}$  was the most efficient route for the removal of the tether. Reaction of **III-12a** in this optimized conditions led to the triol building block **III-15** in an excellent yield of 99%. The removal of the tether in the bis-fullerene derivative **III-13a** gave two products, the mono-adduct derivative **III-16** and the bis-adduct derivative **III-17**.



**Scheme III-2.** Reagents and conditions: (i)  $\text{BF}_3 \cdot \text{Et}_2\text{O}$ ,  $\text{CH}_2\text{Cl}_2/\text{MeCN}$  2:1, rt, 12 h (**III-15**: 99%; **III-16**: 77% and **III-17**: 62%).

As for compound **III-12a**, the Bingel reactions were highly regioselective and an equatorial addition pattern was assigned for the bis-adduct **III-17**. In reason of the in/out stereoisomerism, the unambiguous assignment of the addition pattern was not possible based on the  $C_1$  molecular symmetry deduced from the NMR spectra. The UV/vis spectrum of **III-17** was then recorded and clearly revealed the diagnostic features previously reported for analogous *equatorial*  $C_{60}$  bis-adducts.

In this work, we have shown that the spacer between the malonate subunits and the central *t*-butylsilane group plays an important role in the outcome of the reaction as it acts as a part of the directing tether. The removal of the silyl group in the *e,e,e* tris-adduct **III-12a** and its by-products **III-13a** gave access to racemic polyols derivatives **III-15** and **III-17** with *e* addition patterns. Based on the reactivity of the alcohol functions and on the *e* addition pattern, **III-15** and **III-17** are ideal building blocks for the preparation of mixed  $C_{60}$  hexa-adducts with an octahedral addition pattern.

It should be noted that this work has been done before the publication of a similar work of Hirsch and coworkers in which they used a phosphate tether.<sup>[19]</sup>

### 3. Synthesis of optically pure [60]fullerene *e,e,e*-tris-adducts.

#### 3.1 Introduction.

Fullerene chemistry has quickly generated unprecedented stereochemical problems and the covalent functionalization of fullerenes has produced a plethora of chiral compounds.<sup>[20]</sup> Among them, derivatives with inherently chiral addition pattern, i.e. derivatives of an achiral parent fullerene in which the derivatization creates a chiral functionalization pattern on the fullerene scaffold irrespective of the nature of the addends, are fascinating molecules.<sup>[21]</sup> C<sub>60</sub> multi-adducts with rotational symmetry are typical examples of this family of compounds. A specific nomenclature has been set to describe this type of chirality (**Chapter I-2.3.1**).<sup>[22]</sup>

The C<sub>3</sub>-symmetrical addition pattern *e,e,e* is inherently chiral. As shown in [Figure III-12](#), the achiral parent bis-adduct *e* presents two possible bonds with *e* relationship toward the two already bond addends. Functionalization of the “<sup>f</sup>Re” face leads to the <sup>f</sup>A-tris-adduct *e,e,e*. In opposition, functionalization of the “<sup>f</sup>Si” face leads to <sup>f</sup>C-tris-adducts *e,e,e*. There is no energetic difference between the two double bonds and thus an enantiomeric pair of the <sup>f</sup>A (50%) and <sup>f</sup>C (50%) *e,e,e* tris-adducts is obtained.

Separation of this two enantiomers requires preparative chromatography (HPLC) with a chiral stationary phase.<sup>[23]</sup> An alternative of chiral HPLC is to use chiral addends in order to generate diastereoisomeric compounds. Stepwise additions of chiral addends also required preparative HPLC but with achiral stationary phase.<sup>[24,25]</sup> However separation of the enantiomeric *e,e,e* addition pattern was achieved as diastereoisomeric *e,e,e* tris-adducts.<sup>[24,25]</sup>

Since then, a few strategies allowing the regioselective preparation of racemic fullerene tris(malonates) have been reported but only one example of tris-cyclopropanation of C<sub>60</sub> has been applied in the preparation of enantiomerically pure tris-adducts of C<sub>60</sub> with an *e,e,e* addition pattern.<sup>[26]</sup> In this particular case, the reaction of C<sub>60</sub> with an optically pure macrocyclic tris-malonate derivative provided two separable diastereoisomers with opposite inherently chiral *e,e,e* addition patterns. The formation of the tris-adducts was diastereoselective and a diastereoisomeric excess (de value) of 55% was measured. A major drawback of this strategy is related to the particularly difficult synthesis of the cyclo-tris(malonate) precursor.

An efficient synthetic approach for the preparation of optically pure C<sub>3</sub>-symmetrical tris-adducts still remains an important challenge in fullerene chemistry. In particular, an easy access to the water-soluble tris(malonic acid) derivative of C<sub>60</sub> with an *e,e,e* addition pattern is highly desirable in the context of its remarkable biological activity.<sup>[27]</sup> In this context, we now show that our methodology for the regioselective formation of *e,e,e* tris-adducts of C<sub>60</sub> is perfectly suited for an expeditious synthesis and separation of both <sup>f</sup>A and <sup>f</sup>C enantiomers.

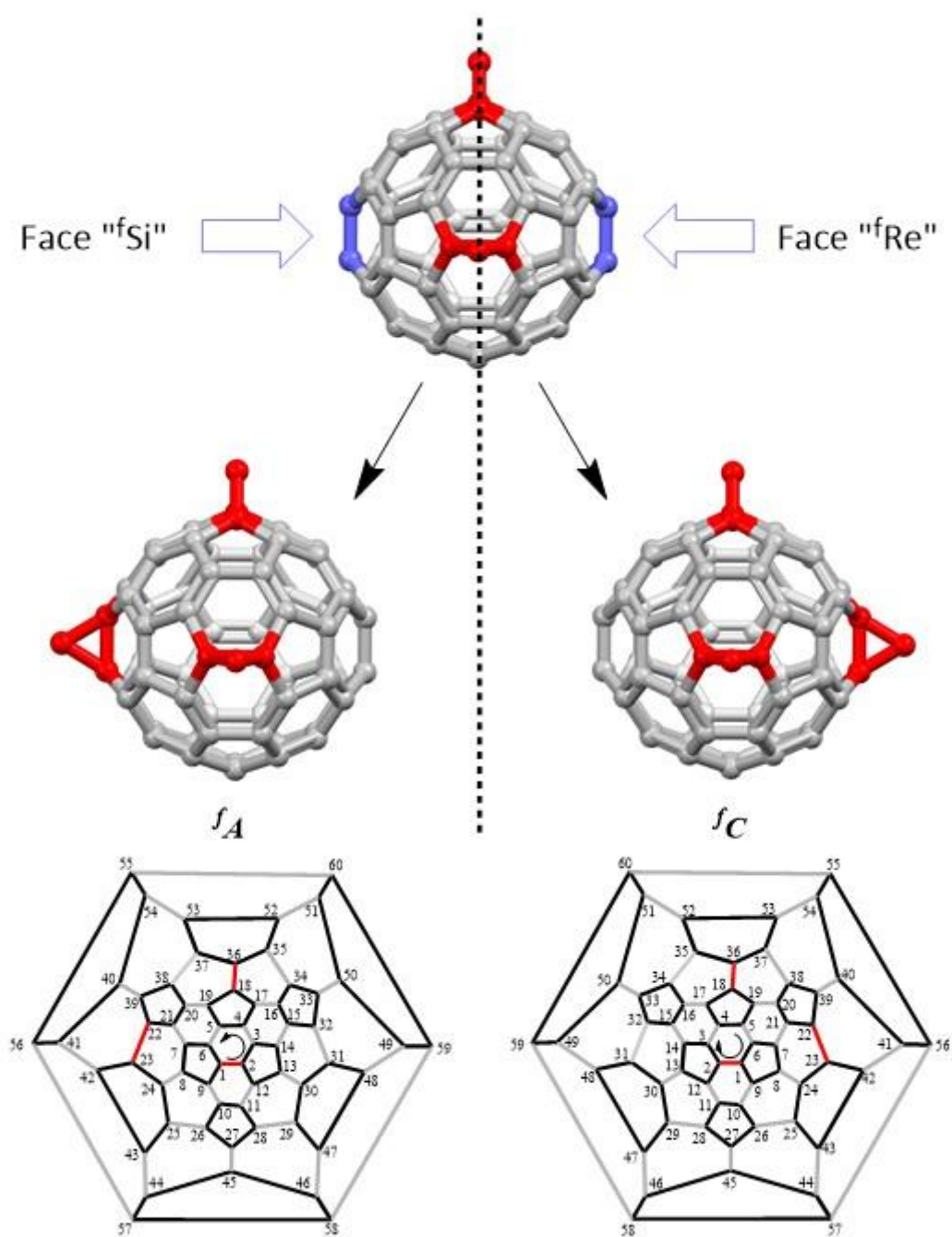
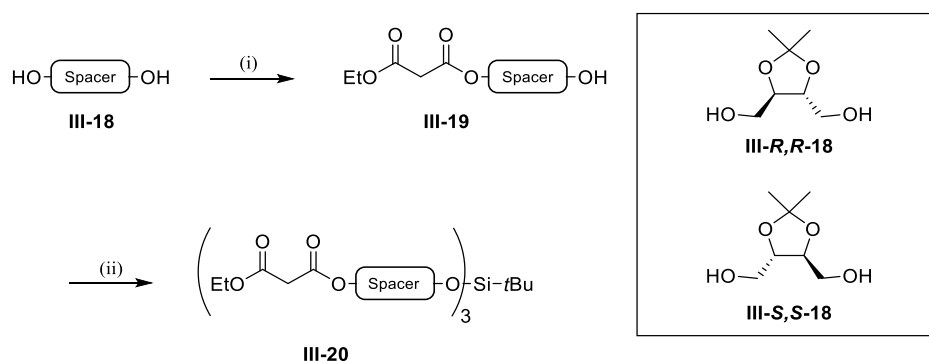


Figure III-12 : Illustration of the chirality of the *e,e,e* addition pattern.

## 3.2 Syntheses.

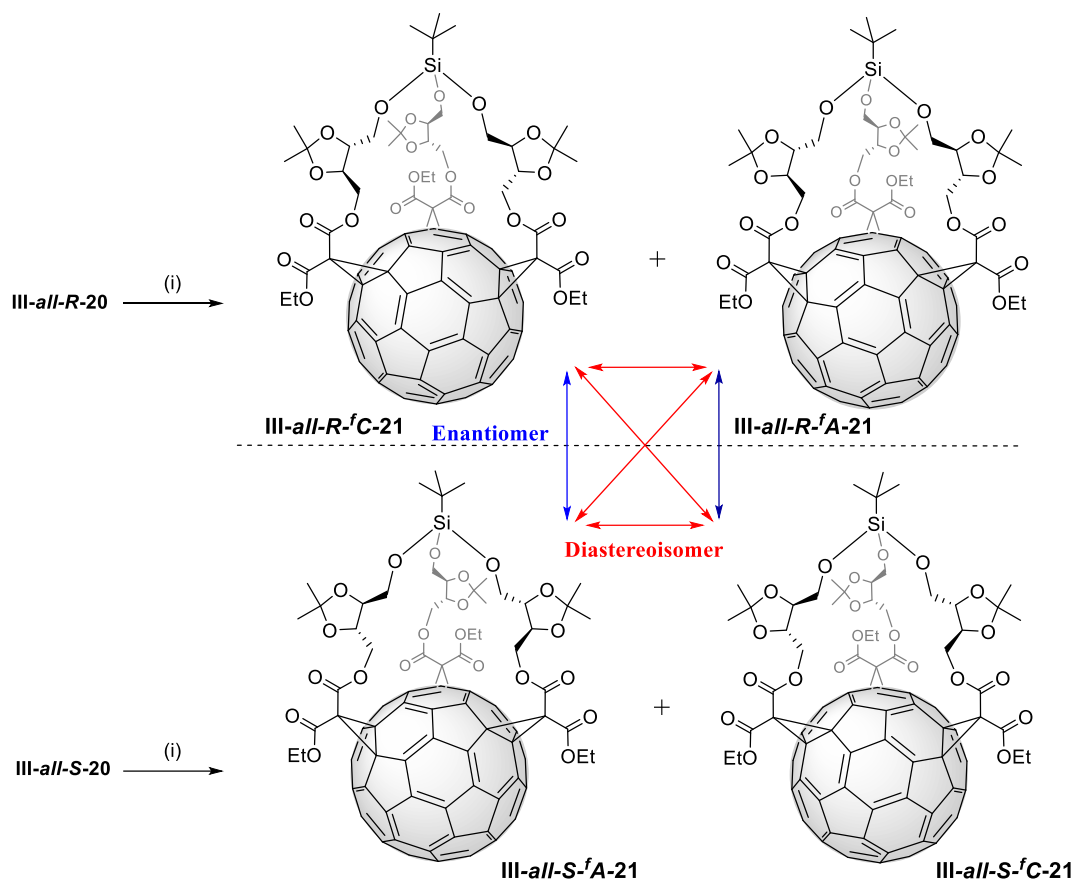
This work has been done in collaboration with Dr. Sebastiano Guerra.

The optically pure Si-tethered tris(malonate) precursors **III-all-R-20** and **III-all-S-20** were prepared from the commercially available  $C_2$ -symmetrical diols **III-R,R-18** and **III-S,S-18**, respectively (**Scheme III-3**). Mono-esterification of diols **III-R,R-18** and **III-S,S-18** with ethyl malonyl chloride followed by reaction of the resulting alcohol with  $t\text{BuSiCl}_3$  in the presence of imidazole gave tris-malonates **III-all-R-20** and **III-all-S-20**, respectively.



**Scheme III-3.** Reagents and conditions: (i) Ethyl malonyl chloride, pyridine, THF, rt, 2 h; (ii)  $t\text{BuSiCl}_3$ , imidazole, DMF, rt, 12 h.

The reaction of both enantiomers of **III-20** with  $C_{60}$  was then performed under the conditions developed for preparation of fullerene  $e,e,e$  tris-adducts from  $t$ -butyl(trialkoxy)silane derivatives (**Scheme III-4**). Specifically, 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU) was added to a solution of **III-all-R-20** or **III-all-S-20**,  $C_{60}$  and iodine in toluene at  $-15^\circ\text{C}$ . Starting from **III-all-R-20**, diastereoisomeric  $e,e,e$  fullerene tris-adducts **III-all-R<sup>f</sup>C-21** and **III-all-R<sup>f</sup>A-21** were obtained. Importantly, a significant difference in polarity between the two diastereoisomers allowed for an easy separation using classical silica column chromatography. The two compounds differ by the absolute configuration of the inherently chiral addition pattern on the fullerene core (<sup>f</sup>C or <sup>f</sup>A,  $f$  = fullerene, C = clockwise and A = anticlockwise). The corresponding enantiomers, **III-all-S<sup>f</sup>A-21** and **III-all-S<sup>f</sup>C-21**, were prepared from **III-all-S-20**. The addition of the Si-tethered tris(malonate) proceeded with complete regioselectivity, giving rise to products with the  $e,e,e$  fullerene addition pattern exclusively. On the other hand, no significant diastereoselectivity was evidenced as the two possible diastereoisomers were thus obtained in an almost 1:1 ratio. Diastereoselectivity was not particularly expected due to the rather flexible structures of the starting  $t$ -butyl(trialkoxy)silane precursors. Similar observations have been also reported by Diederich and co-workers for the addition of optically pure cyclotrimeratrylene-tethered tris(malonate) to  $C_{60}$  leading to almost equal amounts of the two possible diastereoisomeric  $trans$ -3, $trans$ -3 tris-adducts.<sup>[10]</sup>



**Scheme III-4.** Reagents and conditions: (i)  $C_{60}$ ,  $I_2$ , DBU, PhMe,  $-15^\circ\text{C}$ , 1 h (**III-all-R-<sup>f</sup>C-21**: 9% and **III-all-R-<sup>f</sup>A-21**: 9%; **III-all-S-<sup>f</sup>A-21**: 7% and **III-all-S-<sup>f</sup>C-21**: 5%).

For each pair of enantiomers of **III-21**, the relative position of the three addends on the  $C_{60}$  core was determined based on the  $C_3$ -symmetry deduced from their  $^1\text{H}$ - and  $^{13}\text{C}$ -NMR spectra. Furthermore, the four compounds exhibit the classical cherry-red colour of  $e,e,e$  fullerene tris-adducts in solution. Hence, their UV/vis spectra present effectively the characteristic features reported for related fullerene  $e,e,e$  tris-adducts. A tentative assignment of the absolute configuration of the  $e,e,e$  addition pattern on the fullerene sphere in **III-all-R-<sup>f</sup>C-21**, **III-all-R-<sup>f</sup>A-21**, **III-all-S-<sup>f</sup>A-21** and **III-all-S-<sup>f</sup>C-21** was based on the comparison of their circular dichroism (CD) spectra with known literature data. For related  $e,e,e$  tris-adducts, the determination of the absolute configuration was achieved by the comparison of experimental CD spectra with calculated ones for specific optical isomers.<sup>[24]</sup> As shown in [Figure III-13](#), the CD spectra of each pair of enantiomers (**III-all-R-<sup>f</sup>C-21** and **III-all-S-<sup>f</sup>A-21**; **III-all-R-<sup>f</sup>A-21** and **III-all-S-<sup>f</sup>C-21**) display the expected mirror-image shapes with band positions in full agreement with those reported by Hirsch and co-workers for  $e,e,e$  tris-adducts.<sup>[24]</sup> The pronounced Cotton effects observed for these compounds result essentially from the strong chiroptical contributions of the chirally functionalized fullerene chromophore and influence of the chiral tether are minor.<sup>[24,28–31]</sup> Indeed, the CD spectra recorded for compounds **III-all-R-<sup>f</sup>C-21**, **III-all-R-<sup>f</sup>A-21**, **III-all-S-<sup>f</sup>A-21** and **III-all-S-<sup>f</sup>C-21** were not only clear fingerprints allowing the

assignment of their absolute configurations, they also confirmed the  $e,e,e$  addition pattern on the fullerene sphere.

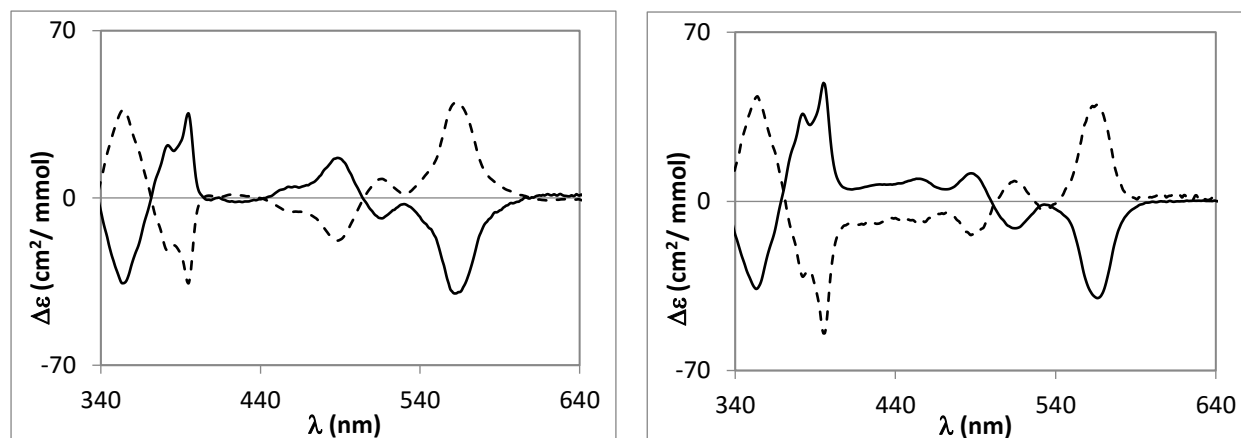


Figure III-13 : Left: CD spectra ( $CH_2Cl_2$ ) of **III-all- $R^fC$ -21** (—) and **III-all- $S^fA$ -21** (- - -); Right: CD spectra ( $CH_2Cl_2$ ) of **III-all- $R^fA$ -21** (- - -) and **III-all- $S^fC$ -21** (—).

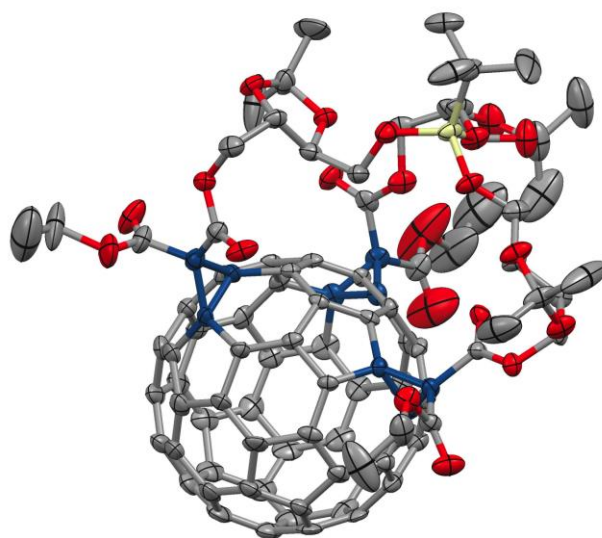
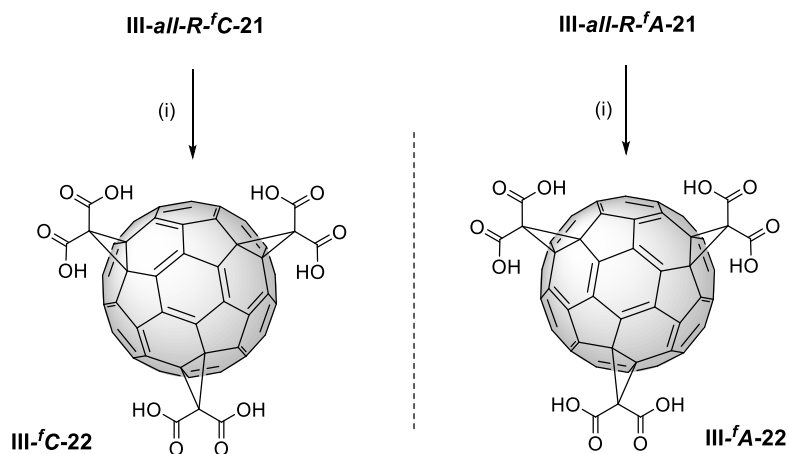


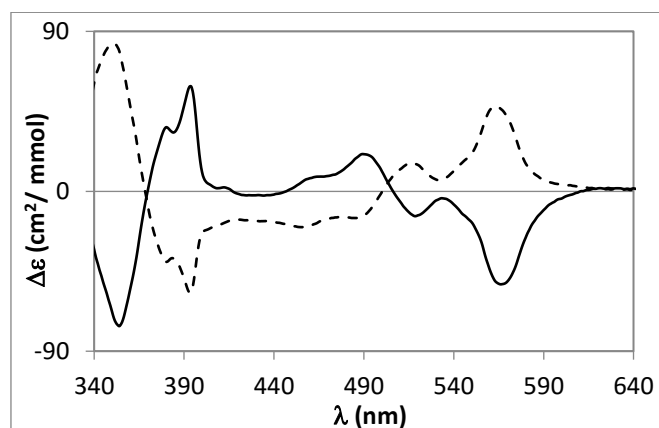
Figure III-14 : ORTEP plot of the structure of **III-all- $S^fA$ -21**.( $CH_2Cl_2$ )<sub>3</sub> (thermal ellipsoids are shown at 50% probability level; for clarity, the co-crystallized solvent molecules are not shown; C: gray; O: red, Si: yellow; H: white). To highlight the chiral  $e,e,e^fA$  addition pattern on the fullerene scaffold, the C atoms of the three cyclopropane rings are shown in dark blue.

For compound **III-all- $S^fA$ -21**, crystals suitable for X-ray crystal structure analysis were obtained by slow diffusion of hexane into a  $CH_2Cl_2$  solution of **III-all- $S^fA$ -21** (Figure III-14). Importantly, the X-ray crystal analysis of **III-all- $S^fA$ -21** unambiguously confirmed the structural assignment based on the CD data, the addition pattern on the fullerene sphere being effectively  $e,e,e$  with an absolute  $^fA$  configuration.

Finally, treatment of diastereoisomers **III-all- $R^f$ C-21** and **III-all- $R^f$ A-21** with NaH/MeOH/PhMe afforded the optically pure tris(malonic acid) derivatives of  $C_{60}$ , **III- $f$ C-22** and **III- $f$ A-22** (Fig. 3). The CD spectra of **III- $f$ C-22** and **III- $f$ A-22** recorded in MeOH are depicted in Figure III-15.



**Scheme III-5.** Reagents and conditions: (i) PhMe, MeOH, NaH, 60 °C (**III- $f$ C-22**: 77%; **III- $f$ A-22**: 87%).



**Figure III-15:** CD spectra (MeOH) of **III- $f$ C-22** (—) and **III- $f$ A-22** (- - -).

## 4. Conclusion.

The expeditious preparation of e,e,e tris-adducts of  $C_{60}$  has been achieved. The key step is a threefold Bingel reaction between  $C_{60}$  and *t*-butyl(trialkoxy)silane derivatives bearing three malonate substituents. The *t*-BuSi group connecting the three reactive malonates acts as a directing unit and the tris-addition is highly regioselective. A major advantage of using a silane-based connecting moiety is related to the easy cleavage of Si-O bonds upon the cyclisation step. The resulting polyols derivatives are valuable building blocks for further chemical transformation based on the reactivity of the alcohol functions. Furthermore, with



their e or e,e,e fullerene addition pattern, these compounds are also perfectly suited for the preparation of fullerene hexa-adducts with an octahedral addition pattern.

Our methodology was also perfectly suited for the preparation of optically pure e,e,e tris-adducts. The reaction of readily available optically active Si-tethered tris(malonates) with C<sub>60</sub> gave easily separable diastereoisomers differing by the absolute configuration of the inherently chiral addition pattern on the fullerene core. Importantly, the absolute configuration of the inherently chiral e,e,e addition pattern has been unambiguously determined using X-ray crystal structure analysis for the first time. Finally, ester hydrolysis of the diastereoisomeric cyclic fullerene derivatives gave easy access to optically pure tris(malonic acid) derivatives of C<sub>60</sub>.

## 5. Bibliography.

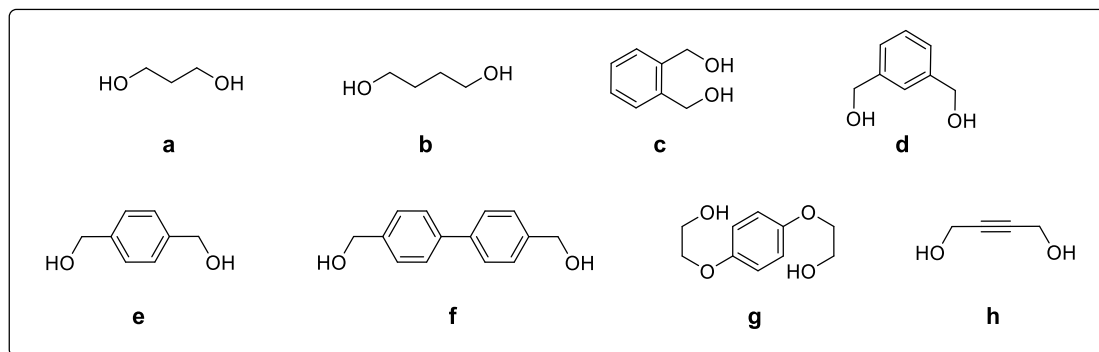
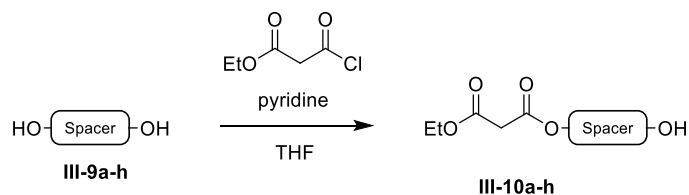
- [1] L. Isaacs, R. F. Haldimann, F. Diederich, *Angew. Chem. Int. Ed. Engl.* **1994**, *33*, 2339–2342.
- [2] C. Thilgen, S. Sergeyev, F. Diederich, in *Templates Chem. I* (Eds.: C.A. Schalley, F. Vögtle, K.H. Dötz), Springer Berlin Heidelberg, **2004**, pp. 1–61.
- [3] J.-F. Nierengarten, T. Habicher, R. Kessinger, F. Cardullo, F. Diederich, V. Gramlich, J.-P. Gisselbrecht, C. Boudon, M. Gross, *Helv. Chim. Acta* **1997**, *80*, 2238–2276.
- [4] J.-P. Bourgeois, L. Echegoyen, M. Fibbioli, E. Pretsch, F. Diederich, *Angew. Chem. Int. Ed.* **1998**, *37*, 2118–2121.
- [5] E. Dietel, A. Hirsch, E. Eichhorn, A. Rieker, S. Hackbarth, B. Röder, *Chem. Commun.* **1998**, 1981–1982.
- [6] S. Sergeyev, F. Diederich, *Angew. Chem. Int. Ed.* **2004**, *43*, 1738–1740.
- [7] R. Kessinger, C. Thilgen, T. Mordasini, F. Diederich, *Helv. Chim. Acta* **2000**, *83*, 3069–3096.
- [8] G. Rapenne, F. Diederich, J. Crassous, A. Collet, L. Echegoyen, *Chem. Commun.* **1999**, 1121–1122.
- [9] J. W. Steed, P. C. Junk, J. L. Atwood, M. J. Barnes, C. L. Raston, R. S. Burkhalter, *J. Am. Chem. Soc.* **1994**, *116*, 10346–10347.
- [10] A. Kraszewska, P. Rivera-Fuentes, G. Rapenne, J. Crassous, A. G. Petrovic, J. L. Alonso-Gómez, E. Huerta, F. Diederich, C. Thilgen, *Eur. J. Org. Chem.* **2010**, *2010*, 4402–4411.
- [11] A. Hirsch, I. Lamparth, H. R. Karfunkel, *Angew. Chem. Int. Ed. Engl.* **1994**, *33*, 437–438.
- [12] F. Beuerle, N. Chronakis, A. Hirsch, *Chem. Commun.* **2005**, 3676–3678.
- [13] F. Beuerle, A. Hirsch, *Chem. – Eur. J.* **2009**, *15*, 7434–7446.
- [14] W. Qian, Y. Rubin, *Angew. Chem. Int. Ed.* **1999**, *38*, 2356–2360.
- [15] R. Schwenninger, T. Müller, B. Kräutler, *J. Am. Chem. Soc.* **1997**, *119*, 9317–9318.
- [16] S.-C. Chuang, S. I. Khan, Y. Rubin, *Org. Lett.* **2006**, *8*, 6075–6078.

- [17] D. Sigwalt, *Conception, Synthèse et Étude de Dérivés de C<sub>60</sub> Fonctionnalisés; Applications Biologiques et Développement Méthodologique.*, Thèse, Université De Strasbourg, **2013**.
- [18] L. E. Echegoyen, F. D. Djojo, A. Hirsch, L. Echegoyen, *J. Org. Chem.* **2000**, *65*, 4994–5000.
- [19] A. Gmehling, W. Donaubaue, F. Hampel, F. W. Heinemann, A. Hirsch, *Angew. Chem. Int. Ed.* **2013**, *52*, 3521–3524.
- [20] C. Thilgen, I. Gosse, F. Diederich, in *Top. Stereochem.* (Ed.: S.E. Denmark), John Wiley & Sons, Inc., **2002**, pp. 1–124.
- [21] C. Thilgen, F. Diederich, *Chem. Rev.* **2006**, *106*, 5049–5135.
- [22] W. H. Powell, F. Cozzi, G. P. Moss, C. Thilgen, R. J.-R. Hwu, A. Yerin, *Pure Appl. Chem.* **2002**, *74*, 629–695.
- [23] B. Gross, V. Schurig, I. Lamparth, A. Herzog, F. Djojo, A. Hirsch, *Chem. Commun.* **1997**, 1117–1118.
- [24] F. Djojo, A. Hirsch, *Chem. – Eur. J.* **1998**, *4*, 344–356.
- [25] F. Djojo, A. Hirsch, S. Grimme, *Eur. J. Org. Chem.* **1999**, *1999*, 3027–3039.
- [26] N. Chronakis, A. Hirsch, *Chem. Commun.* **2005**, 3709–3711.
- [27] L. L. Dugan, D. M. Turetsky, C. Du, D. Lobner, M. Wheeler, C. R. Almlı, C. K.-F. Shen, T.-Y. Luh, D. W. Choi, T.-S. Lin, *Proc. Natl. Acad. Sci.* **1997**, *94*, 9434–9439.
- [28] M. Maggini, G. Scorrano, A. Bianco, C. Toniolo, M. Prato, *Tetrahedron Lett.* **1995**, *36*, 2845–2846.
- [29] H. Goto, N. Harada, J. Crassous, F. Diederich, *J. Chem. Soc. Perkin Trans. 2* **1998**, 1719–1724.
- [30] F. Giacalone, J. L. Segura, N. Martín, *J. Org. Chem.* **2002**, *67*, 3529–3532.
- [31] S. Campidelli, P. Bourgun, B. Guintchin, J. Furrer, H. Stoeckli-Evans, I. M. Saez, J. W. Goodby, R. Deschenaux, *J. Am. Chem. Soc.* **2010**, *132*, 3574–3581.
- [32] I. Lamparth, A. Hirsch, *J. Chem. Soc. Chem. Commun.* **1994**, 1727–1728.

## 6. Experimental part.

**General:** Reagents and solvents were purchased as reagent grade and used without further purification. All reactions were performed in standard glassware under an inert Ar atmosphere. Evaporation and concentration were done at water aspirator pressure and drying in vacuo at 10<sup>-2</sup> Torr. Column chromatography: silica gel 60 (230-400 mesh, 0.040-0.063 mm) was purchased from E. Merck. Thin Layer Chromatography (TLC) was performed on aluminum sheets coated with silica gel 60 F<sub>254</sub> purchased from E. Merck. IR spectra (cm<sup>-1</sup>) were recorded on a Perkin-Elmer Spectrum One Spectrophotometer. UV/Visible spectra were recorded on a Perkin-Elmer Lambda 35 UV/Vis Spectrometer. NMR spectra were recorded on a Bruker AC 300 or AC 400 with solvent peaks as reference at 25°C. MALDI-TOF-MS were obtained on a Bruker ULTRAFLEX TOF/TOF mass spectrometer with a dithranol matrix.

## Synthesis.



**General Procedures. Preparation of III-10a-h (GP I):** A solution of ethyl malonyl chloride (1 equiv.) in anhydrous THF was added dropwise within 30 min. to a solution of appropriate diol (III-9a-h, 2-4 equiv.) and pyridine (2 equiv.) in anhydrous THF at 0 °C. The mixture was then stirred at room temperature overnight. The resulting mixture was filtered on  $\text{SiO}_2$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 8:2 to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 97:3) gave III-10a-h.

**Compound III-10a:** III-10a was synthesized according to GP I from diol III-9a (1.1 mL, 15 mmol), ethyl malonyl chloride (1.0 mL, 7.5 mmol) and pyridine (1.1 mL, 15 mmol) in anhydrous THF (75 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 8:2 to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 97:3) yielded III-10a (1.07 g, 5.6 mmol, 74 %) as a colorless oil.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta$  = 4.36 (t,  $^3J$  = 7 Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{OOC}$ ), 4.23 (q,  $^3J$  = 7 Hz, 2 H,  $\text{CH}_3\text{CH}_2$ ), 3.75 (q,  $^3J$  = 6 Hz, 2 H,  $\text{HOCH}_2\text{CH}_2$ ), 3.41 (s, 2 H,  $\text{OOCCH}_2\text{COO}$ ), 1.92 (quint.,  $^3J$  = 6 Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{CH}_2$ ), 1.87 (t,  $^3J$  = 6 Hz, 1 H,  $\text{HOCH}_2$ ), 1.31 (t,  $^3J$  = 7 Hz, 3 H,  $\text{CH}_3\text{CH}_2$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 75 MHz):  $\delta$  = 166.9, 166.7, 62.7, 61.7, 59.2, 41.6, 31.5, 14.1 ppm.

**Compound III-10b:** III-10b was synthesized according to GP I from diol III-9b (5.4 mL, 60 mmol), ethyl malonyl chloride (1.9 mL, 15 mmol) and pyridine (2.3 mL, 30 mmol) in anhydrous THF (130 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 97:3) yielded III-10b (2.52 g, 12.3 mmol, 82 %) as a colorless oil. IR (neat):  $\nu$  = 3442 (br, OH), 1728 (C=O)  $\text{cm}^{-1}$ .  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta$  = 4.23 (q,  $^3J$  = 7 Hz, 2 H,  $\text{CH}_3\text{CH}_2$ ), 4.22 (t,  $^3J$  = 6 Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{OOC}$ ), 3.70 (q,  $^3J$  = 6 Hz, 2 H,  $\text{HOCH}_2\text{CH}_2$ ), 3.39 (s, 2 H,  $\text{OOCCH}_2\text{COO}$ ), 1.83-1.65 (m, 4 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{CH}_2$ ), 1.45 (m, 1 H,  $\text{HOCH}_2$ ), 1.30 (t,  $^3J$  = 7 Hz, 3 H,  $\text{CH}_3\text{CH}_2$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 75 MHz):  $\delta$  = 166.6, 65.3, 62.3, 61.6, 41.7, 29.0, 24.9, 14.1 ppm.

**Compound III-10c:** III-10c was synthesized according to GP I from diol III-9c (3.0 g, 21.7 mmol), ethyl malonyl chloride (1.4 mL, 10.8 mmol) and pyridine (1.7 mL, 21.7 mmol) in anhydrous THF (100 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 97:3) yielded III-10c (1.30 g, 5.1 mmol, 47 %) as a colorless oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz): δ = 7.48-7.28 (m, 4 H, ArH), 5.33 (s, 2 H, ArCH<sub>2</sub>OOC), 4.78 (s, 2 H, ArCH<sub>2</sub>OH), 4.20 (q, <sup>3</sup>J = 7 Hz, 2 H, CH<sub>3</sub>CH<sub>2</sub>), 3.43 (s, 2 H, OOCCH<sub>2</sub>COO), 2.18 (br s, 1 H, HOCH<sub>2</sub>), 1.26 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>3</sub>CH<sub>2</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz): δ = 166.6, 166.3, 139.5, 133.1, 130.1, 129.2, 129.1, 128.2, 65.2, 62.9, 61.8, 41.6, 14.0 ppm.

**Compound III-10d:** III-10d was synthesized according to GP I from diol III-9d (3.0 g, 21.7 mmol), ethyl malonyl chloride (1.8 mL, 14 mmol) and pyridine (2.3 mL, 28.5 mmol) in anhydrous THF (100 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded III-10d (1.03 g, 4.1 mmol, 28 %) as a colorless oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz): δ = 7.36 (m, 4 H, ArH), 5.19 (s, 2 H, ArCH<sub>2</sub>OOC), 4.71 (s, 2 H, ArCH<sub>2</sub>OH), 4.20 (q, <sup>3</sup>J = 7 Hz, 2 H, CH<sub>3</sub>CH<sub>2</sub>), 3.42 (s, 2 H, OOCCH<sub>2</sub>COO), 1.64 (br s, 1H, HOCH<sub>2</sub>), 1.25 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>3</sub>CH<sub>2</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz): δ = 166.5, 141.8, 135.6, 128.8, 127.4, 127.0, 126.7, 67.1, 64.9, 61.7, 41.6, 14.0 ppm.

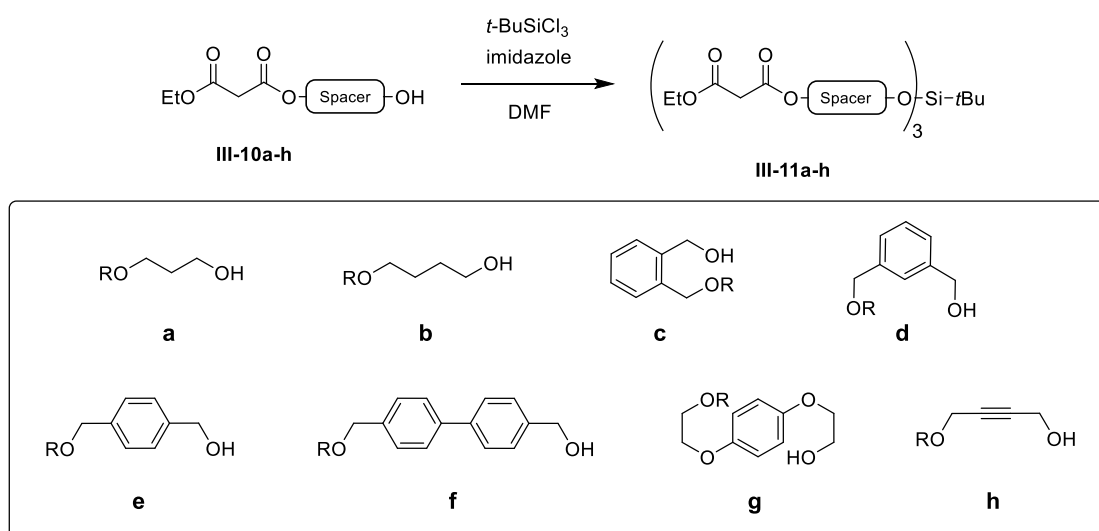
**Compound III-10e:** III-10e was synthesized according to GP I from diol III-9e (5.0 g, 36.2 mmol), ethyl malonyl chloride (2.3 mL, 18.1 mmol) and pyridine (2.8 mL, 34.6 mmol) in anhydrous THF (300 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded III-10e (2.08 g, 8.2 mmol, 45 %) as a colorless oil. IR(neat): ν = 3442 (br, OH), 1727 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz): δ = 7.38 (m, 4 H, ArH), 5.20 (s, 2 H, ArCH<sub>2</sub>OOC), 4.72 (d, <sup>3</sup>J = 5 Hz, 2 H, ArCH<sub>2</sub>OH), 4.21 (q, <sup>3</sup>J = 7 Hz, 2H, CH<sub>3</sub>CH<sub>2</sub>), 3.43 (s, 2H, OOCCH<sub>2</sub>COO), 1.76 (br s, 1H, HOCH<sub>2</sub>), 1.27 (t, <sup>3</sup>J = 7 Hz, 3H, CH<sub>3</sub>CH<sub>2</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz): δ = 166.5, 166.4, 141.2, 134.7, 128.6, 127.1, 66.9, 65.0, 61.6, 41.6, 14.0 ppm.

**Compound III-10f:** III-10f was synthesized according to GP I from diol III-9f (2.5 g, 11.6 mmol), ethyl malonyl chloride (0.8 mL, 5.8 mmol) and imidazole (0.79 g, 11.6 mmol) in anhydrous THF (150 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded III-10f (0.82 g, 2.5 mmol, 43 %) as a colorless oil. IR(neat): ν = 3454 (br, OH), 1732 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz): δ = 7.59 (2d, <sup>3</sup>J = 8 Hz, 2 H, ArH), 7.58 (2d, <sup>3</sup>J = 8 Hz, 2 H, ArH), 7.45 (2d, <sup>3</sup>J = 8 Hz, 2 H, ArH), 7.44 (2d, <sup>3</sup>J = 8 Hz, 2 H, ArH), 5.23 (s, 2 H, ArCH<sub>2</sub>OOC), 4.75 (s, 2 H, ArCH<sub>2</sub>OH), 4.20 (q, <sup>3</sup>J = 7 Hz, 2 H, CH<sub>3</sub>CH<sub>2</sub>), 3.44 (s, 2 H, OOCCH<sub>2</sub>COO), 1.26 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>3</sub>CH<sub>2</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz): δ = 166.5, 166.4, 141.0, 140.2, 140.0, 134.4, 128.9, 127.5, 127.3, 127.2, 66.9, 65.1, 61.6, 41.7, 14.1 ppm.

**Compound III-10g:** III-10g was synthesized according to GP I from diol III-9g (2.5 g, 12.6 mmol), ethyl malonyl chloride (.74 mL, 4.2 mmol) and imidazole (0.57 g, 8.4 mmol) in anhydrous THF (110 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded III-10g (2.35 g, 7.5 mmol, 65 %) as a colorless oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): δ = 6.85 (s, 4 H, ArH),

4.48 (t,  $^3J = 5$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{OOC}$ ), 4.19 (q,  $^3J = 7$  Hz, 2 H,  $\text{COOCH}_2\text{CH}_3$ ), 4.14 (t,  $^3J = 5$  Hz, 2 H,  $\text{OCH}_2\text{CH}_2\text{OOC}$ ), 4.03 (t,  $^3J = 5$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{OH}$ ), 3.93 (t,  $^3J = 5$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{OH}$ ), 3.41 (s, 2 H,  $\text{OOCCH}_2\text{COO}$ ), 1.26 (t,  $^3J = 7$  Hz, 3 H,  $\text{CH}_2\text{CH}_3$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 166.6$ , 166.4, 153.2, 152.9, 115.8, 115.6, 69.8, 66.4, 63.8, 61.6 (two peaks), 41.5, 14.0 ppm.

**Compound III-10h:** III-10h was synthesized according to GP I from diol III-9h (3.8 g, 43.9 mmol), ethyl malonyl chloride (2.9 mL, 22.6 mmol) and pyridine (3.4 mL, 43.9 mmol) in anhydrous THF (130 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 8:2 to  $\text{CH}_2\text{Cl}_2$ ) yielded III-10h (2.71 g, 13.5 mmol, 59 %) as a colorless oil.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 4.78$  (t,  $^4J = 2$  Hz, 2 H,  $\text{COOCH}_2\text{C}\equiv\text{CCH}_2$ ), 4.30 (t,  $^4J = 2$  Hz, 2 H,  $\text{CH}_2\text{C}\equiv\text{CCH}_2\text{OH}$ ), 4.21 (q,  $^3J = 7$  Hz, 2 H,  $\text{CH}_3\text{CH}_2$ ), 3.41 (s, 2 H,  $\text{OOCCH}_2\text{COO}$ ), 1.81 (s, 1 H, OH), 1.28 (t,  $^3J = 7$  Hz, 3 H,  $\text{CH}_3\text{CH}_2$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 75 MHz):  $\delta = 166.1$ , 165.9, 85.5, 79.1, 61.7, 53.2, 51.0, 41.3, 14.0 ppm.



**General Procedures. Preparation of III-11a-h (GP II):** A mixture of  $t\text{BuSiCl}_3$  (1 equiv.), the appropriate alcohol (III-10a-h; 3.2 equiv.) and imidazole (3.5 equiv.) in anhydrous DMF was stirred at room temperature overnight. Then, water was added and the aqueous layer extracted with  $\text{Et}_2\text{O}$ . The organic layer was washed, dried ( $\text{MgSO}_4$ ) and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 97:3) gave III-11a-h.

**Compound III-11a:** III-11a was synthesized according to GP II from III-10a (1.06 g, 5.6 mmol),  $t\text{BuSiCl}_3$  (0.34 g, 1.8 mmol) and imidazole (0.43 g, 6.3 mmol) in anhydrous DMF (11 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 97:3) yielded III-11a (0.76 g, 1.2 mmol, 66 %) as a colorless oil.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 4.28$  (t,  $^3J = 7$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{OOC}$ ), 4.20 (q,  $^3J = 7$  Hz, 6 H,  $\text{CH}_3\text{CH}_2$ ), 3.89 (t,  $^3J = 6$  Hz, 6 H,  $\text{SiOCH}_2\text{CH}_2$ ), 3.38 (s, 6 H,  $\text{OOCCH}_2\text{COO}$ ), 1.92 (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2$ ), 1.29 (t,  $^3J = 7$  Hz, 9 H,  $\text{CH}_3\text{CH}_2$ ), 0.98 (s, 9 H,  $\text{Si}(\text{CH}_3)_3$ )

ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz):  $\delta$  = 166.6, 166.5, 62.4, 61.5, 59.5, 41.6, 31.5, 26.3, 17.7, 14.1 ppm.

**Compound III-11b:** III-11b was synthesized according to GP II from III-10b (1.00 g, 4.9 mmol), *t*BuSiCl<sub>3</sub> (0.29 g, 1.5 mmol) and imidazole (0.36 g, 5.3 mmol) in anhydrous DMF (10 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 97:3) yielded III-11b (0.66 g, 1.0 mmol, 63 %) as a colorless oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta$  = 4.22 (t, <sup>3</sup>J = 7 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.20 (q, <sup>3</sup>J = 7 Hz, 6 H, CH<sub>3</sub>CH<sub>2</sub>), 3.81 (t, <sup>3</sup>J = 6 Hz, 6 H, SiOCH<sub>2</sub>CH<sub>2</sub>), 3.39 (s, 6 H, OOCCH<sub>2</sub>COO), 1.62-1.59 (m, 12 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.30 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 0.96 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz):  $\delta$  = 166.7, 166.6 (6 C, OOCCH<sub>2</sub>COO), 65.3, 62.5, 61.5, 41.6, 28.8, 26.4, 25.0, 17.7, 14.1 ppm.

**Compound III-11c:** III-11c was synthesized according to GP II from III-10c (1.00 g, 4.0 mmol), *t*BuSiCl<sub>3</sub> (0.24 g, 1.2 mmol) and imidazole (0.30 g, 4.3 mmol) in anhydrous DMF (15 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded III-11c (1.04 g, 1.2 mmol, 51 %) as a colorless oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta$  = 7.30 (m, 12 H, ArH), 5.15 (s, 6 H, ArCH<sub>2</sub>OOC), 4.87 (s, 6 H, ArCH<sub>2</sub>OSi), 4.15 (q, <sup>3</sup>J = 7 Hz, 6 H, CH<sub>3</sub>CH<sub>2</sub>), 3.35 (s, 6 H, OOCCH<sub>2</sub>COO), 1.22 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 1.02 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz):  $\delta$  = 166.4, 166.3, 138.6, 132.7, 129.1, 128.6, 127.6, 127.4, 64.5, 62.9, 61.6, 41.5, 26.9, 26.3, 17.9, 14.0 ppm.

**Compound III-11d:** III-11d was synthesized according to GP II from III-10d (1.0 g, 4 mmol), *t*BuSiCl<sub>3</sub> (0.17 g, 0.9 mmol) and imidazole (0.21 g, 3.1 mmol) in anhydrous DMF (15 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded III-11d (0.47 g, 0.6 mmol, 44 %) as a colorless oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta$  = 7.28 (m, 12 H, ArH), 5.14 (s, 6 H, ArCH<sub>2</sub>OOC), 4.87 (s, 6 H, ArCH<sub>2</sub>OSi), 4.17 (q, <sup>3</sup>J = 7 Hz, 6 H, CH<sub>3</sub>CH<sub>2</sub>), 3.38 (s, 6 H, OOCCH<sub>2</sub>COO), 1.23 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 1.06 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz):  $\delta$  = 166.5, 165.4, 140.9, 135.4, 128.3, 127.1, 126.6, 126.3, 67.1, 64.8, 61.6, 41.6, 18.0, 14.0 ppm.

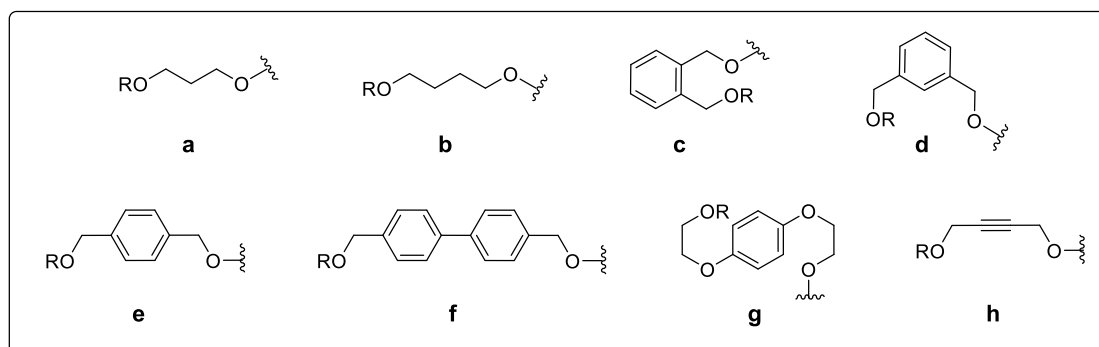
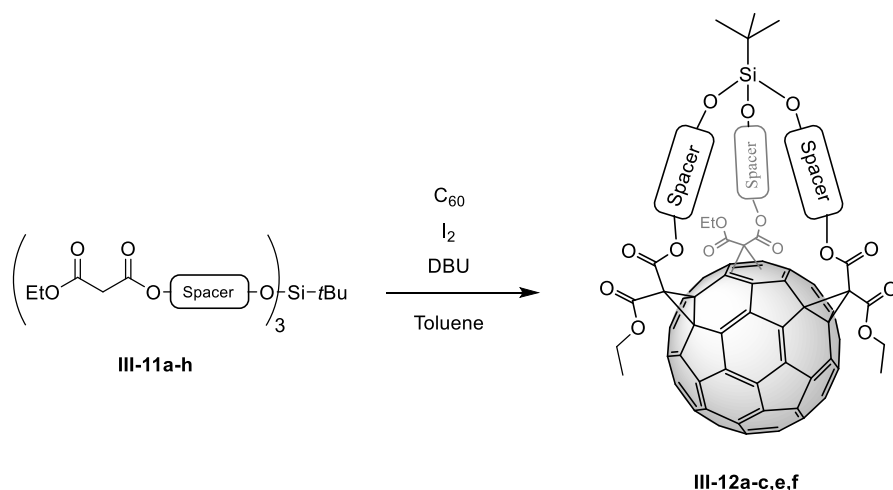
**Compound III-11e:** III-11e was synthesized according to GP II from III-10e (1.5 g, 5.9 mmol), *t*BuSiCl<sub>3</sub> (0.36 g, 1.9 mmol) and imidazole (0.44 g, 6.5 mmol) in anhydrous DMF (12 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded III-11e (0.97 g, 1.2 mmol, 62 %) as a colorless oil. IR(neat):  $\nu$  = 1730 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta$  = 7.33 (m, 12 H, ArH), 5.20 (s, 6 H, ArCH<sub>2</sub>OOC), 4.88 (s, 6 H, ArCH<sub>2</sub>OSi), 4.21 (q, <sup>3</sup>J = 7 Hz, 6 H, CH<sub>3</sub>CH<sub>2</sub>), 3.44 (s, 6 H, OOCCH<sub>2</sub>COO), 1.27 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 1.07 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz):  $\delta$  = 166.5, 166.4, 140.1, 134.2, 128.3, 126.4, 67.0, 64.8, 61.6, 41.6, 26.4, 18.0, 14.0 ppm.

**Compound III-11f:** III-11f was synthesized according to GP II from III-10f (0.82 g, 2.5 mmol), *t*BuSiCl<sub>3</sub> (0.15 g, 0.8 mmol) and imidazole (0.19 g, 2.8 mmol) in anhydrous DMF (10 mL).

Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded **III-11f** (0.48 g, 0.45 mmol, 56 %) as a colorless oil. IR(neat):  $\nu = 1733$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.57$  (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 7.53 (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 7.42 (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 7.39 (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 5.23 (s, 6 H, ArCH<sub>2</sub>OOC), 4.95 (s, 6 H, ArCH<sub>2</sub>OSi), 4.20 (q, <sup>3</sup>J = 7 Hz, 6 H, CH<sub>3</sub>CH<sub>2</sub>), 3.44 (s, 6 H, OOCCH<sub>2</sub>COO), 1.26 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 1.12 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.5, 166.4, 141.2, 139.9, 139.5, 134.3, 128.8, 127.2, 127.0, 126.8, 67.0, 64.9, 61.6, 41.7, 26.5, 18.1, 14.1$  ppm.

**Compound III-11g:** **III-11g** was synthesized according to **GP II** from **III-10g** (1.74 g, 5.5 mmol), *t*BuSiCl<sub>3</sub> (0.34 g, 1.8 mmol) and imidazole (0.43 g, 6.3 mmol) in anhydrous DMF (10 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>) yielded **III-11g** (0.67 g, 0.65 mmol, 37 %) as a colorless oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 6.80$  (s, 12 H, ArH), 4.47 (t, <sup>3</sup>J = 5 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.19 (q, <sup>3</sup>J = 7 Hz, 6 H, COOCH<sub>2</sub>CH<sub>3</sub>), 4.15 (t, <sup>3</sup>J = 5 Hz, 6 H, OCH<sub>2</sub>CH<sub>2</sub>OOC), 4.12 (t, <sup>3</sup>J = 5 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>OSi), 4.01 (t, <sup>3</sup>J = 5 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>OSi), 3.42 (s, 6 H, OOCCH<sub>2</sub>COO), 1.26 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>2</sub>CH<sub>3</sub>), 1.00 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.6, 166.4, 153.5, 152.6, 115.7, 115.6, 69.6, 66.4, 63.8, 62.1, 61.6, 41.5, 26.2, 17.8, 14.0$  ppm.

**Compound III-11h:** **III-11h** was synthesized according to **GP II** from **III-10h** (1.72 g, 8.6 mmol), *t*BuSiCl<sub>3</sub> (0.52 g, 2.7 mmol) and imidazole (0.64 g, 9.3 mmol) in anhydrous DMF (20 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 97:3) yielded **III-11h** (0.79 g, 1.2 mmol, 43 %) as a colorless oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta = 4.79$  (t, <sup>4</sup>J = 2 Hz, 6 H, COOCH<sub>2</sub>C≡CCH<sub>2</sub>), 4.52 (t, <sup>4</sup>J = 2 Hz, 6 H, CH<sub>2</sub>C≡CCH<sub>2</sub>OSi), 4.20 (q, <sup>3</sup>J = 7 Hz, 6 H, CH<sub>3</sub>CH<sub>2</sub>), 3.41 (s, 6 H, OOCCH<sub>2</sub>COO), 1.28 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>3</sub>CH<sub>2</sub>), 0.97 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz):  $\delta = 166.1, 165.9, 85.1, 78.6, 61.7, 53.2, 51.9, 41.2, 25.8, 17.5, 14.0$  ppm.



**General Procedures. Preparation of *e,e,e*-Trisadducts III-12a-h (GP III):** DBU (7.5 equiv.) was added to a solution of  $C_{60}$  (1.5 equiv.), the appropriate tris-malonate III-11a-h (1 equiv.) and  $I_2$  (3.5 equiv.) in toluene (2 mL / mg of  $C_{60}$ ) at  $-15^\circ\text{C}$ . The resulting mixture was stirred for 1 h at  $-15^\circ\text{C}$ , then filtered on  $\text{SiO}_2$  (cyclohexane to  $\text{CH}_2\text{Cl}_2$ ) and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 1:1) followed by gel permeation chromatography (Biobeads SX-1,  $\text{CH}_2\text{Cl}_2$ ) or by recrystallization by slow diffusion in THF/cyclohexane if necessary gave III-12a-c,e,f.

**Compound III-12a & III-13a:** III-12a & III-13a were synthesized according to GP III from III-11a (153 mg, 0.23 mmol),  $C_{60}$  (250 mg, 0.35 mmol),  $I_2$  (213 mg, 0.84 mmol) and DBU (0.26 mL, 1.7 mmol) in toluene (500 mL). Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 5:5) followed by gel permeation chromatography (Biobeads SX-1,  $\text{CH}_2\text{Cl}_2$ ) afforded III-12a (86 mg, 0.06 mmol, 26 %) as a cherry-red glassy solid and III-13a (53 mg, 0.03 mmol, 11%) as a brown glassy solid. III-12a: IR(neat):  $\nu = 1740$  (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 230$  (102500), 251 (102000), 280 (67000), 302 (sh, 54100), 381 (sh, 6100), 484 (4500), 566 (sh, 1200) nm.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 4.54$  (m, 3 H), 4.41 (m, 6 H,  $\text{CH}_3\text{CH}_2$ ), 4.22 (m, 3 H), 3.59 (t,  $^3J = 6$  Hz, 6 H,  $\text{SiOCH}_2\text{CH}_2$ ), 1.93 (m, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2$ ), 1.38 (t,  $^3J = 7$  Hz, 9 H,  $\text{CH}_3\text{CH}_2$ ), 0.89 (s, 9 H,  $\text{Si}(\text{CH}_3)_3$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 163.6, 163.3, 147.2, 147.0, 146.9, 146.7, 146.5, 146.4$  (two peaks), 146.3, 145.9, 144.3, 143.6, 143.3, 143.2, 143.1 (two peaks), 142.8, 141.8, 141.0, 70.9, 70.2, 63.9, 63.4, 59.8, 31.7, 26.1, 17.5, 14.2 ppm. MALDI-TOF-MS: 1366.2 ( $[\text{M}]^+$ , calcd. for  $\text{C}_{88}\text{H}_{42}\text{O}_{15}\text{Si}$ : 1366.2). III-13a: IR(neat):  $\nu = 1740$  (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 392$



(sh, 11100), 408 (sh, 6900), 426 (5600), 481 (5300), 547 (sh, 3000), 612 (sh, 1200), 689 (300) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz): 4.89-4.81 (m, 1H), 4.77-4.70 (m, 1H), 4.65 (t, <sup>3</sup>J = 6 Hz, 2H), 4.59 (q, <sup>3</sup>J = 7 Hz, 2H), 4.56-4.48 (m, 4H), 4.46-4.38 (m, 1H), 4.33-4.26 (m, 1H), 4.07 (t, <sup>3</sup>J = 6 Hz, 2H), 3.91-3.73 (m, 4H), 2.16 (quint., <sup>3</sup>J = 6 Hz, 2H), 2.08-1.97 (m, 4H), 1.51 (t, <sup>3</sup>J = 7 Hz, 3H), 1.45 (m, 6H), 0.93 (s, 9H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz): 163.6 (two peaks), 163.5, 163.4, 163.1, 148.6, 147.3, 147.2, 146.8, 146.6, 146.5, 146.4, 146.3, 146.1, 146.0, 145.7, 145.5, 145.3 (two peaks), 145.2 (two peaks), 145.1, 145.0, 144.9 (two peaks), 144.8, 144.7 (two peaks), 144.6 (two peaks), 144.5, 144.3, 144.1, 144.0, 143.9, 143.8, 143.7 (two peaks), 143.6 (two peaks), 143.5, 143.4, 143.3, 143.1, 143.0, 142.9, 142.5, 142.4, 142.2, 142.0, 141.9, 141.8, 141.7, 141.6, 141.5, 141.3, 141.0, 140.6, 139.8, 139.7, 139.0, 138.9, 138.6, 71.8, 71.7 (two peaks), 71.6, 70.5, 64.4, 63.6, 63.5, 63.4, 63.1, 60.0, 59.7, 58.6, 53.9, 52.3, 51.6, 32.1, 31.7, 31.5, 26.3, 17.6, 14.3, 14.2 ppm. MALDI-TOF-MS: 2087.2 ([M]<sup>+</sup>, calcd for C<sub>148</sub>H<sub>42</sub>O<sub>15</sub>Si: 2087.2).

**Compound III-12b:** III-12b was synthesized according to **GP III** from III-11b (159 mg, 0.23 mmol), C<sub>60</sub> (250 mg, 0.35 mmol), I<sub>2</sub> (213 mg, 0.84 mmol) and DBU (0.26 mL, 1.7 mmol) in toluene (500 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5) followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) afforded III-12b (44 mg, 0.03 mmol, 14%) as a cherry-red glassy solid. IR(neat):  $\nu = 1747$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 252$  (74900), 281 (50600), 305 (sh, 38600), 380 (3900), 484 (3300), 564 (sh, 960) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.53$  (m, 3 H), 4.43 (m, 6 H, CH<sub>3</sub>CH<sub>2</sub>), 4.09 (m, 3 H), 3.64 (m, 6 H, SiOCH<sub>2</sub>CH<sub>2</sub>), 1.73 (quint., <sup>3</sup>J = 7 Hz, 6 H), 1.51 (m, 6 H), 1.39 (t, <sup>3</sup>J = 8 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 0.93 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.4$ , 163.2, 146.9 (two peaks), 146.8, 146.7 (two peaks), 146.5, 146.3 (two peaks), 145.8, 144.2 (two peaks), 143.7, 143.3, 143.0, 142.6, 142.3, 142.1, 141.0, 71.0, 70.1, 66.4, 63.3, 62.2, 43.5, 30.2, 28.2, 27.0, 26.4, 25.2, 17.9, 14.2 ppm. MALDI-TOF-MS: 1408.2 ([M]<sup>+</sup>, calcd. for C<sub>91</sub>H<sub>48</sub>O<sub>15</sub>Si: 1408.3).

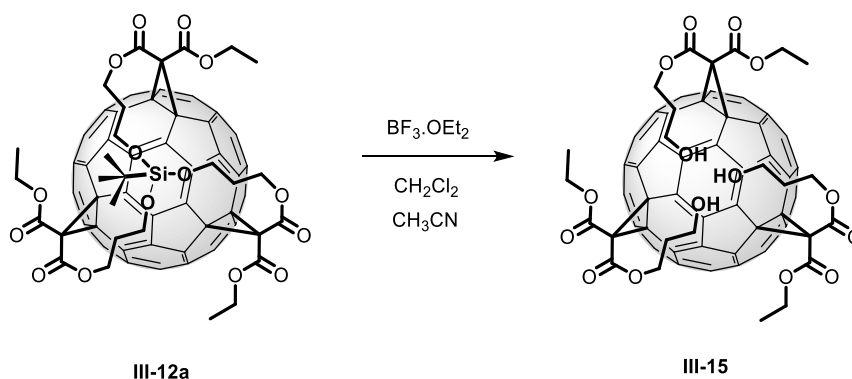
**Compound III-12c:** III-12c was synthesized according to **GP III** from III-11c (388 mg, 0.46 mmol), C<sub>60</sub> (500 mg, 0.69 mmol), I<sub>2</sub> (408 mg, 1.6 mmol) and DBU (0.54 mL, 3.4 mmol) in toluene (1 L). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 1:1) followed by recrystallization by slow diffusion in THF/cyclohexane afforded III-12c (165 mg, 0.11 mmol, 23%) as a cherry-red glassy solid. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 374$  (11800), 468 (5700), 557 (sh, 1600) nm. <sup>1</sup>H NMR (300 MHz, CDCl<sub>3</sub>):  $\delta = 7.34$ -7.22 (m, 12 H, ArH), 5.42 (d, 3J = 11 Hz, 3 H), 5.26 (d, 3J = 11 Hz, 3 H), 4.78 (d, 3J = 11 Hz, 3 H), 4.60 (d, J = 11 Hz, 3 H), 4.36 (m, 6 H), 1.33 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 0.86 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (75 MHz, CDCl<sub>3</sub>):  $\delta = 164.3$ , 163.2, 147.2, 146.8, 146.5 (two peaks), 146.3, 146.2 (two peaks), 145.7, 145.6, 144.4, 144.2, 144.1, 143.2, 142.3, 141.8, 141.0, 140.7, 139.4, 132.0, 130.8, 130.1, 127.9, 127.8, 70.3, 69.7, 66.3, 63.2, 62.5, 52.4, 26.1, 17.6, 14.3 ppm. MALDI-TOF-MS: 1554.3 ([M+H]<sup>+</sup>, calcd for C<sub>103</sub>H<sub>49</sub>O<sub>15</sub>Si: 1554.3).

**Compound III-12e:** III-12e was synthesized according to **GP III** from III-11e (262 mg, 0.31 mmol), C<sub>60</sub> (340 mg, 0.47 mmol), I<sub>2</sub> (279 mg, 1.1 mmol) and DBU (0.35 mL, 2.3 mmol) in toluene (675 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 1:1) followed by

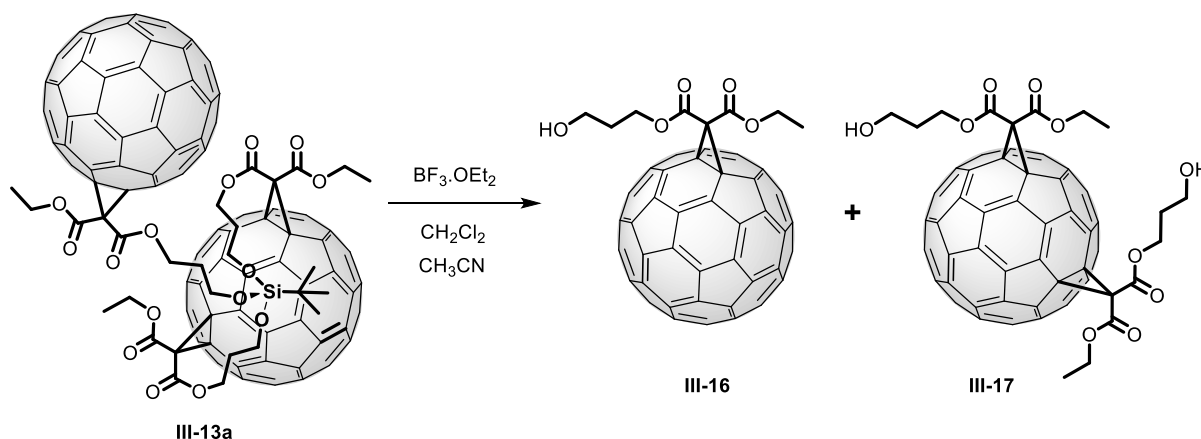
recrystallization by slow diffusion in THF/cyclohexane afforded **III-12e** (48 mg, 0.03 mmol, 10%) as a cherry-red glassy solid. IR(neat):  $\nu = 1746$  (C=O)  $\text{cm}^{-1}$ . UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 230$  (236200), 249 (229700), 283 (153200), 305 (sh, 120600), 383 (2700), 479 (1500), 563 (sh, 500) nm. <sup>1</sup>H NMR (300 MHz, CDCl<sub>3</sub>):  $\delta = 7.03$  (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 6.81 (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 5.44 (d, J = 13 Hz, 3 H), 5.06 (d, J = 13 Hz, 3 H), 4.42 (d, J = 12 Hz, 3 H), 4.35 (q, <sup>3</sup>J = 7 Hz, 6 H), 4.16 (d, J = 12 Hz, 3 H), 1.30 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 0.98 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (75 MHz, CDCl<sub>3</sub>):  $\delta = 163.6, 163.3, 146.9, 146.8, 146.7, 146.5$  (two peaks), 146.4, 146.3, 145.8, 144.5, 144.4, 144.1, 143.3, 143.0, 142.4, 142.3, 142.2, 141.0, 140.7 (two peaks), 133.9, 128.1, 126.5, 70.8, 70.1, 68.2, 65.3, 63.4, 53.0, 26.9, 26.2, 18.1, 14.2 ppm. MALDI-TOF-MS: 1553.3 ([M]<sup>+</sup>, calcd for C<sub>103</sub>H<sub>48</sub>O<sub>15</sub>Si: 1553.3).

**Compound III-14e (all-trans 3): III-14e (all-trans 3)** was synthesized according to **GP III** from **III-11e** (1.59 g, 1.89 mmol), C<sub>60</sub> (1.50 g, 2.07 mmol), I<sub>2</sub> (1.68 g, 6.6 mmol) and DBU (2.22 mL, 14.1 mmol) in toluene (3 L). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 1:1) afforded **III-14e (all-trans 3)** (30 mg, 0.02 mmol, 1%) as a purple-red glassy solid. IR(neat):  $\nu = 1745$  (C=O)  $\text{cm}^{-1}$ . UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 246$  (42200), 302 (19300), 407 (sh, 1600), 493 (1500), 574 (sh, 760) nm. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.32$  (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 7.10 (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 5.88 (d, J = 11 Hz, 3 H), 4.94 (q, <sup>3</sup>J = 7 Hz, 6 H), 4.91 (d, J = 11 Hz, 3 H), 4.40 (m, 6 H), 1.37 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 1.04 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 163.6, 148.3, 147.6, 147.1, 146.2, 146.0, 145.5, 145.4, 145.1, 145.0, 143.7, 143.0, 142.9, 142.7, 142.0, 141.8, 141.6, 141.2, 138.8, 133.5, 131.2, 124.7, 71.2, 70.9, 68.8, 64.6, 63.4, 53.4, 50.4, 26.4, 19.3, 14.2$  ppm. MALDI-TOF-MS: 1553.3 ([M]<sup>+</sup>, calcd for C<sub>103</sub>H<sub>48</sub>O<sub>15</sub>Si: 1553.3).

**Compound III-12f: III-12f** was synthesized according to **GP III** from **III-11f** (480 mg, 0.45 mmol), C<sub>60</sub> (357 mg, 0.49 mmol), I<sub>2</sub> (399 mg, 1.6 mmol) and DBU (0.51 mL, 3.4 mmol) in toluene (720 mL). Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5) followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) afforded **III-12f** (25 mg, 0.01 mmol, 3%) as a cherry-red glassy solid. IR(neat):  $\nu = 1739$  (C=O)  $\text{cm}^{-1}$ . UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 253$  (67800), 279 (sh, 39900), 304 (sh, 22600), 483 (1900), 565 (sh, 600) nm. <sup>1</sup>H NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 7.40$  (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 7.28 (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 7.24 (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 6.86 (d, <sup>3</sup>J = 8 Hz, 6 H, ArH), 5.53 (d, J = 12 Hz, 3 H), 5.05 (d, J = 12 Hz, 3 H), 4.63 (d, J = 12 Hz, 3 H), 4.51 (d, J = 12 Hz, 3 H), 4.39 (m, 6 H), 1.33 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>3</sub>CH<sub>2</sub>), 1.06 (s, 9 H, SiC(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 164.1, 163.4, 148.0, 146.9, 146.7$  (two peaks), 146.6, 146.5 (two peaks), 146.3, 145.9, 145.6, 145.1, 144.3, 143.5, 143.3, 141.7, 141.3, 141.1, 141.0, 140.7, 139.9, 139.6, 133.8, 128.5, 127.9, 127.7, 127.0, 70.9, 69.9, 68.9, 65.6, 63.2, 52.7, 26.9, 18.6, 14.2 ppm. MALDI-TOF-MS: 1781.5 (100%, [M]<sup>+</sup>, calcd for C<sub>121</sub>H<sub>60</sub>O<sub>15</sub>Si: 1781.3), 1736.5 (7%, [M-OEt]<sup>+</sup>, calcd for C<sub>119</sub>H<sub>55</sub>O<sub>14</sub>Si: 1736.3).

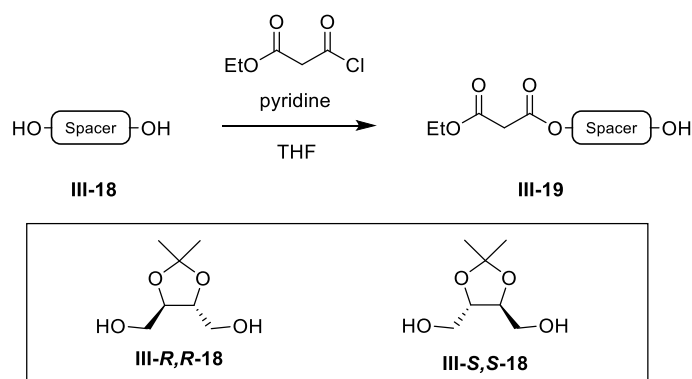


**Compound III-15:**  $\text{BF}_3 \cdot \text{Et}_2\text{O}$  (0.32 mL, 1.8 mmol) was added to a solution of **III-12a** (252 mg, 0.18 mmol) in  $\text{CH}_2\text{Cl}_2/\text{CH}_3\text{CN}$  (4:2 mL) at room temperature. The resulting mixture was stirred overnight at room temperature. A saturated  $\text{NaHCO}_3$  aqueous solution was added and the aqueous layer extracted with  $\text{CH}_2\text{Cl}_2$ . The organic layer was washed with water, dried ( $\text{MgSO}_4$ ) and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{MeOH}$  96:4) afforded **III-15** (229 mg, 0.18 mmol, 99%) as a cherry-red glassy solid. IR(neat):  $\nu = 3377$  (br, OH),  $1737$  ( $\text{C}=\text{O}$ )  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 251$  (115000), 282 (80000), 304 (sh, 60600), 353 (sh, 16400), 379 (sh, 7000), 485 (5100), 564 (sh, 1500) nm.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 4.51$ - $4.35$  (m, 12 H), 3.68 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{OH}$ ), 2.23 (br s, 3 H,  $\text{HOCH}_2$ ), 1.94 (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2$ ), 1.38 (t,  $^3J = 7$  Hz, 9 H,  $\text{CH}_3\text{CH}_2$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 75 MHz):  $\delta = 163.6$ , 163.4, 147.1, 146.9, 146.7, 146.6, 146.5, 146.4, 146.3, 145.8, 145.7, 144.8, 144.6, 144.3, 143.4, 142.7, 142.6, 141.9, 141.8, 141.0, 70.8, 70.1, 64.0, 63.3, 59.0, 31.3, 14.2 ppm. MALDI-TOF-MS: 1284.1 ( $[\text{M}]^+$ , calcd for  $\text{C}_{84}\text{H}_{36}\text{O}_{15}$ : 1284.2).



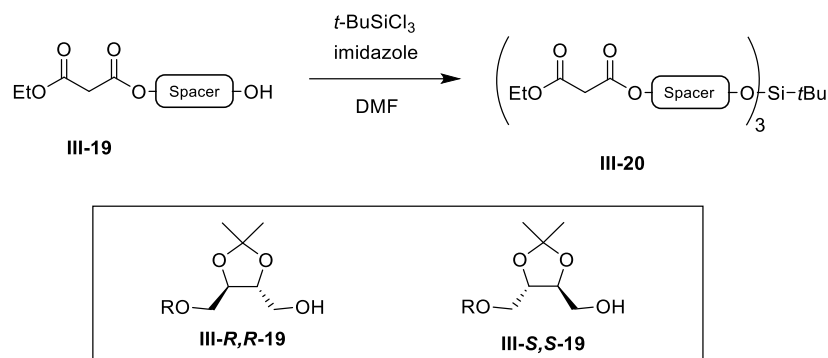
**Compound III-16 & III-17:**  $\text{BF}_3 \cdot \text{Et}_2\text{O}$  (0.32 mL, 2.60 mmol) was added to a solution of **III-13a** (274 mg, 0.13 mmol) in  $\text{CH}_2\text{Cl}_2/\text{CH}_3\text{CN}$  (4:2 mL). The resulting mixture was stirred overnight at room temperature. A saturated  $\text{NaHCO}_3$  aqueous solution was added and the aqueous layer extracted with  $\text{CH}_2\text{Cl}_2$ . The organic layer was washed with water, dried ( $\text{MgSO}_4$ ) and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{MeOH}$  96 : 4) yielded **III-16** (90 mg, 0.10 mmol, 77%) as a brown glassy solid and **III-17** (86 mg, 0.08 mmol, 62%) as red-brown

glassy solid. **III-16**: IR(neat):  $\nu = 3370$  (OH),  $1743$  (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 257$  (106000),  $326$  (33600),  $394$  (sh, 4000),  $400$  (sh, 2900),  $414$  (sh, 2100),  $426$  (2200),  $488$  (1300),  $551$  (sh, 860),  $605$  (sh, 470),  $688$  (180) nm.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 4.59$  (t,  $^3J = 6$  Hz, 2 H,  $\text{COOCH}_2\text{CH}_2$ ),  $4.50$  (q,  $^3J = 7$  Hz, 2 H,  $\text{COOCH}_2\text{CH}_3$ ),  $3.79$  (t,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{OH}$ ),  $2.04$  (quint.,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{CH}_2$ ),  $1.42$  (t,  $^3J = 7$  Hz, 3 H,  $\text{CH}_2\text{CH}_3$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 163.9$ ,  $163.6$ ,  $145.3$  (two peaks),  $145.2$  (two peaks),  $145.1$ ,  $144.9$ ,  $144.7$ ,  $144.6$ ,  $143.9$ ,  $143.6$ ,  $143.1$ ,  $143.0$  (three peaks),  $142.2$ ,  $141.9$ ,  $141.0$  (two peaks),  $139.1$ ,  $138.9$ ,  $71.6$ ,  $64.3$ ,  $63.6$ ,  $63.4$ ,  $59.1$ ,  $52.2$ ,  $34.2$ ,  $31.6$ ,  $30.3$ ,  $29.7$ ,  $29.5$ ,  $14.3$ ,  $14.2$  ppm. MALDI-TOF-MS:  $908.0$  ( $[\text{M}]^+$ , calcd for  $\text{C}_{68}\text{H}_{12}\text{O}_5$ :  $908.1$ ). **III-17**: IR(neat):  $\nu = 3450$  (br, OH),  $1740$  (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 306$  (sh, 21900),  $331$  (sh, 7400),  $397$  (sh, 2400),  $409$  (sh, 1600),  $421$  (1400),  $478$  (sh, 1600) nm.  $^1\text{H}$  NMR (400 MHz,  $\text{CDCl}_3$ ):  $\delta = 4.61$ - $4.48$  (m, 8 H),  $3.80$  (m, 4 H),  $2.04$  (m, 4 H),  $1.94$  (br s, 2 H, OH),  $1.46$  (t,  $^3J = 7$  Hz, 3 H,  $\text{CH}_2\text{CH}_3$ ),  $1.45$  (t,  $^3J = 7$  Hz, 3 H,  $\text{CH}_2\text{CH}_3$ ) ppm.  $^{13}\text{C}$  NMR (100 MHz,  $\text{CDCl}_3$ ):  $\delta = 163.9$ ,  $163.7$ ,  $163.6$ ,  $163.5$ ,  $147.7$ ,  $147.5$ ,  $147.3$  (two peaks),  $146.5$  (two peaks),  $146.2$ ,  $146.1$ ,  $146.0$  (two peaks),  $145.6$  (two peaks),  $145.4$  (two peaks),  $145.2$  (two peaks),  $145.1$ ,  $144.8$  (two peaks),  $144.7$  (two peaks),  $144.6$  (two peaks),  $144.4$  (two peaks),  $144.3$  (two peaks),  $144.2$  (two peaks),  $144.1$  (two peaks),  $143.8$  (two peaks),  $143.7$  (two peaks),  $143.5$  (two peaks),  $143.2$ ,  $143.0$  (four peaks),  $142.6$ ,  $142.2$ ,  $142.0$ ,  $141.9$ ,  $141.8$ ,  $141.6$  (two peaks),  $141.5$ ,  $138.9$  (two peaks),  $138.7$  (two peaks),  $71.6$ ,  $70.3$ ,  $64.2$  (two peaks),  $63.5$ ,  $63.4$ ,  $59.1$ ,  $53.5$  (two peaks),  $51.3$ ,  $31.5$  (two peaks),  $14.2$  ppm. MALDI-TOF-MS:  $1096.1$  ( $[\text{M}]^+$ , calcd for  $\text{C}_{76}\text{H}_{24}\text{O}_{10}$ :  $1096.1$ ).



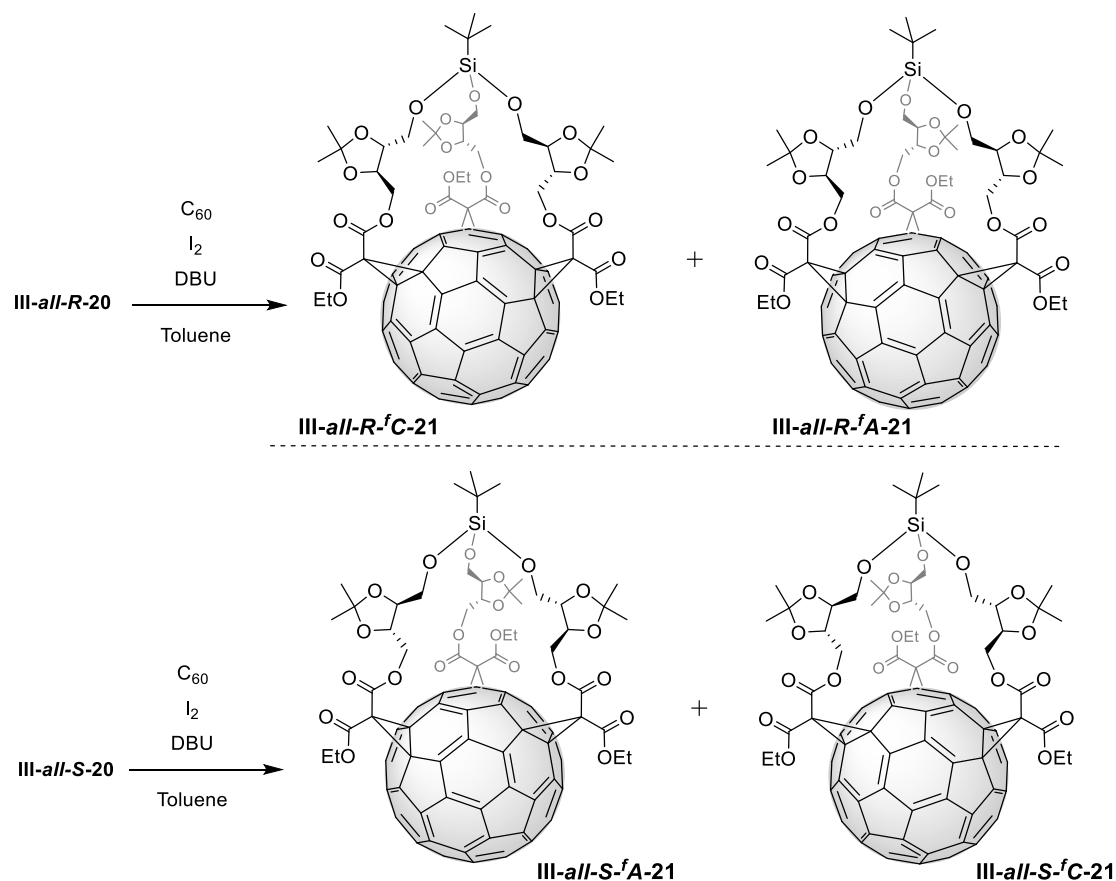
**Compound III-S,S-19**: A solution of ethyl malonyl chloride (0.80 mL, 6.25 mmol) in anhydrous THF (60 mL) was added dropwise over 1 h to a stirred solution of (4*S*,5*S*)-2,2-dimethyl-1,3-dioxolane-4,5-dimethanol (**III-S,S-18**) (2.03 g, 12.50 mmol) and pyridine (0.95 mL, 11.75 mmol) in anhydrous THF (20 mL) at  $0^\circ\text{C}$ . The mixture was then stirred at room temperature overnight. The resulting mixture was filtered to remove the salts and concentrated. Column chromatography ( $\text{SiO}_2$ ,  $\text{CH}_2\text{Cl}_2$ ) gave **III-S,S-19** (1.21 g, 70%) as a colorless oil.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $4.40$ - $4.15$  (m, 5H),  $3.98$  (m, 1H),  $3.86$  (m, 1H),  $3.67$  (m, 1H),  $3.45$  (s, 2H),  $1.93$  (m, 1H),  $1.45$  (s, 3H),  $1.44$  (s, 3H),  $1.31$  (t,  $J = 7$  Hz, 3H).  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 75 MHz):  $166.35$ ,  $166.3$ ,  $109.9$ ,  $78.3$ ,  $74.8$ ,  $64.9$ ,  $61.8$ ,  $61.7$ ,  $41.4$ ,  $27.0$ ,  $26.9$ ,  $14.0$ .

**Compound III-*R,R*-19:** As described for III-*S,S*-19 starting from (4*R*,5*R*)-2,2-dimethyl-1,3-dioxolane-4,5-dimethanol (III-*R,R*-18) (2.00 g, 12.33 mmol), pyridine (1.00 mL, 12.33 mmol) and ethyl malonyl chloride (0.79 mL, 6.17 mmol). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) gave III-*R,R*-19 (1.31 g, 77%) as a colorless oil. <sup>1</sup>H and <sup>13</sup>C-NMR data are rigorously identical to those described for the corresponding enantiomer III-*S,S*-19.



**Compound III-*all-S*-20:** A mixture of III-*S,S*-19 (1.14 g, 4.10 mmol), imidazole (262 mg, 3.85 mmol) and *t*BuSiCl<sub>3</sub> (248 mg, 1.30 mmol) in anhydrous DMF (10 mL) was stirred at 0°C for 1 h. The mixture was then allowed to warm slowly to room temperature and stirred for 12 h at this temperature, then H<sub>2</sub>O (50 mL) was added. The aqueous layer was extracted with Et<sub>2</sub>O (3 x). The combined organic layers were washed with water, dried over MgSO<sub>4</sub>, filtered and concentrated. Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) gave III-*all-S*-20 (460 mg, 39%) as a colorless oil. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz): 4.49-4.42 (m, 3H), 4.27-4.15 (m, 12H), 4.05-3.92 (m, 9H), 3.45 (s, 6H), 1.43 (s, 18H), 1.30 (t, *J* = 7 Hz, 9H), 1.00 (s, 9H). <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz): 166.4, 166.2, 109.9, 65.2, 63.3, 61.6, 41.3, 27.1, 26.9, 26.2, 17.8, 14.0.

**Compound III-*all-R*-20:** As described for III-*all-S*-20 starting from III-*R,R*-19 (1.00 g, 3.62 mmol), imidazole (269 mg, 3.96 mmol) and *t*BuSiCl<sub>3</sub> (217 mg, 1.13 mmol). Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>) gave III-*all-R*-20 (538 mg, 52 %) as a colorless oil. <sup>1</sup>H and <sup>13</sup>C NMR data rigorously identical to those described for the corresponding enantiomer III-*all-S*-20.



**Compound III-all-S<sup>f</sup>A-21 & III-all-S<sup>f</sup>C-21:** DBU (0.56 mL, 3.75 mmol) was added to a stirred solution of  $C_{60}$  (536 mg, 0.75 mmol), III-all-S-20 (460 mg, 0.50 mmol) and  $I_2$  (444 mg, 1.75 mmol) in toluene (1.2 L) at  $-15^\circ\text{C}$ . After 1 h, the mixture was filtered through a short plug of  $\text{SiO}_2$ , eluting first with toluene (to remove unreacted  $C_{60}$ ), then with  $\text{CH}_2\text{Cl}_2$ . Gel permeation chromatography (Biobeads SX-1,  $\text{CH}_2\text{Cl}_2$ ) followed by column chromatography ( $\text{SiO}_2$ , cyclohexane/EtOAc 80:20) gave III-all-S<sup>f</sup>A-21 (59.4 mg, 7%) and III-all-S<sup>f</sup>C-21 (41.4 mg, 5%).

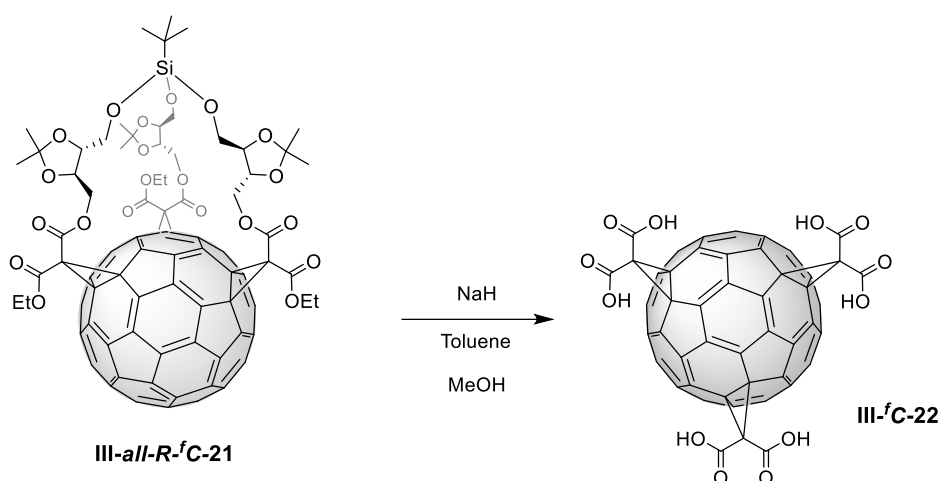
**III-all-S<sup>f</sup>A-21:** Red solid.  $^1\text{H-NMR}$  ( $\text{CDCl}_3$ , 300 MHz): 1.00 (s, 9H), 1.41 (t,  $J = 7$  Hz, 9H), 1.415 (br s, 9H), 1.49 (br s, 9H), 3.69 (d,  $J = 5$  Hz, 6H), 3.92 (td,  $J = 8$  Hz and 5 Hz, 3H), 4.09 (br d,  $J = 8$  Hz, 3H), 4.35-4.51 (m, 9H), 4.57 (dd,  $J = 13$  Hz and 2 Hz, 3H).  $^{13}\text{C-NMR}$  ( $\text{CDCl}_3$ , 100 MHz): 14.1, 18.0, 26.5, 26.7, 27.2, 52.8, 62.2, 63.4, 63.8, 70.1, 70.8, 75.5, 75.7, 109.7, 141.7, 141.7, 142.0, 142.6, 143.0, 143.3, 143.5, 144.1, 144.2, 145.8, 146.2, 146.25, 146.3, 146.4, 146.8, 146.9, 146.9, 147.0, 162.6, 162.7. IR (neat): 1750 (C=O). MALDI-TOF-MS: 1647.31 ( $[\text{M}+\text{Na}]^+$ , calcd for  $\text{C}_{100}\text{H}_{60}\text{O}_{21}\text{SiNa}$ : 1647.33), 1624.33 ( $[\text{M}]^+$ , calcd for  $\text{C}_{100}\text{H}_{60}\text{O}_{21}\text{Si}$ : 1624.34).

**III-all-S<sup>f</sup>C-21:** Red solid. IR (neat): 1749 (C=O).  $^1\text{H-NMR}$  ( $\text{CDCl}_3$ , 300 MHz): 1.05 (s, 9H), 1.40 (t,  $J = 7$  Hz, 9H), 1.42 (s, 18H), 3.69 (dd,  $J = 11$  Hz and 2 Hz, 3H), 3.81-3.90 (m, 6H), 4.30-4.50 (m, 15H).  $^{13}\text{C-NMR}$  ( $\text{CDCl}_3$ , 75 MHz): 14.0, 18.4, 26.8, 26.9, 27.3, 52.8, 62.5, 63.3, 68.2, 70.0, 70.7, 75.3, 78.1, 110.1, 141.0, 141.8, 142.2, 142.8, 143.3 (2C), 143.4, 143.7, 144.3, 145.7, 146.3, 146.4 (2C), 146.5, 146.7, 146.85 (2C), 146.9, 163.0, 163.6. MALDI-TOF-MS: 1647.3 ( $[\text{M}+\text{Na}]^+$ , calcd for  $\text{C}_{100}\text{H}_{60}\text{O}_{21}\text{SiNa}$ : 1647.33), 1624.3 ( $[\text{M}]^+$ , calcd for  $\text{C}_{100}\text{H}_{60}\text{O}_{21}\text{Si}$ : 1624.34).

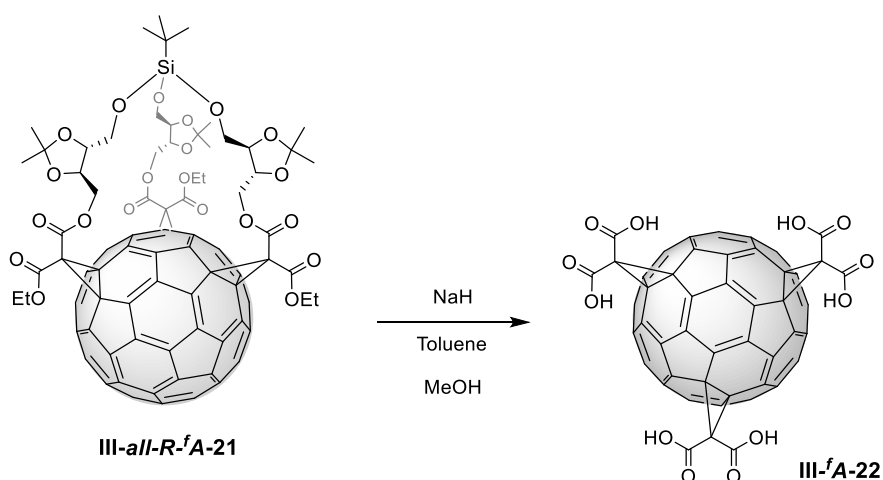
**Compound III-*all-R*<sup>f</sup>A-21 & III-*all-R*<sup>f</sup>C-21:** As described for III-*all-S*<sup>f</sup>A-21 and III-*all-S*<sup>f</sup>C-21 starting from  $C_{60}$  (468 mg, 0.65 mmol), III-*all-R*-20 (538 mg, 0.59 mmol) and  $I_2$  (524 mg, 2.07 mmol). Gel permeation chromatography (Biobeads SX-1,  $CH_2Cl_2$ ) followed by column chromatography ( $SiO_2$ , cyclohexane/EtOAc 80:20) gave III-*all-R*<sup>f</sup>C-21 (80.1 mg, 9%) and III-*all-R*<sup>f</sup>A-21 (81.6 mg, 9%).

*Data for III-*all-R*<sup>f</sup>C-21.* Red solid.  $^1H$  and  $^{13}C$  NMR, IR, MS and UV/vis data rigorously identical to those described for the corresponding enantiomer III-*all-S*<sup>f</sup>A-21.

*Data for III-*all-R*<sup>f</sup>A-21.* Red solid.  $^1H$  and  $^{13}C$  NMR, IR, MS and UV/vis data rigorously identical to those described for the corresponding enantiomer III-*all-S*<sup>f</sup>C-21.



**Compound III-<sup>f</sup>C-22:** A solution of III-*all-R*<sup>f</sup>C-21 (70.3 mg, 0.04 mmol) and NaH (60% dispersion in mineral oil, 19.2 mg, 0.80 mmol) in toluene (10 mL) was stirred at 60°C for 3 h. Then, MeOH (1 mL) was added. After 1 h, the resulting precipitate was collected by centrifugation, washed with toluene, an aqueous 2M  $H_2SO_4$  solution and water. Finally, drying *in vacuo* at 60°C overnight gave III-<sup>f</sup>C-22 (31.5 mg, 77 %) as a red solid. The analytical data of III-<sup>f</sup>C-22 were in complete agreement with literature data.<sup>[32]</sup>



**Compound III-*f*A-22:** A solution of **III-*all-R-f*A-21** (75.3 mg, 0.046 mmol) and NaH (60% dispersion in mineral oil, 22.2 mg, 0.93 mmol) in toluene (10 mL) was stirred at 60°C for 3 h. Then, MeOH (1 mL) was added. After 1 h, the resulting precipitate was collected by centrifugation, washed with toluene, an aqueous 2M H<sub>2</sub>SO<sub>4</sub> solution and water. Finally, drying *in vacuo* at 60°C overnight gave **III-*f*A-22** (39.3 mg, 87%) as a red solid. The analytical data of **III-*f*A-22** were in complete agreement with literature data.<sup>[32]</sup>





**Chapter IV : Syntheses of multi-functionalized hexa-adducts by sequential “click” reactions.**

## 1. Introduction.

The geometry of  $T_h$ -symmetrical  $C_{60}$  hexa-adducts with an octahedral addition pattern is unique in organic chemistry (Figure IV-1). Each addend has four *e* neighbors and a fifth addend in a *trans-1* relationship. This addition pattern can also be compared to an octahedral motif.

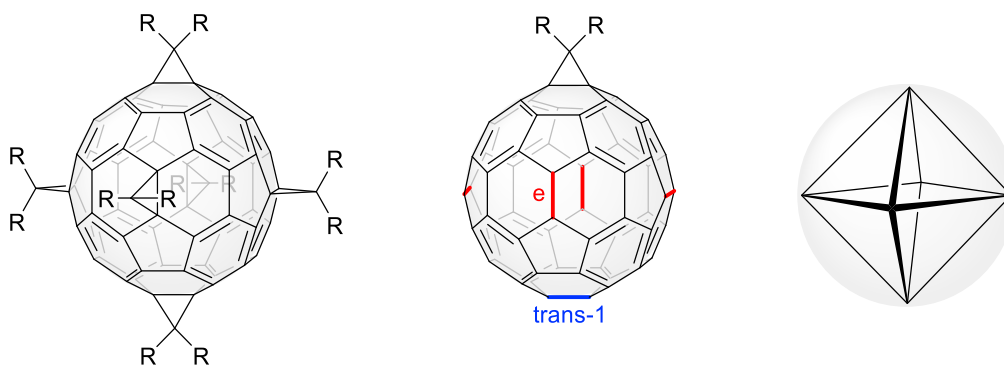
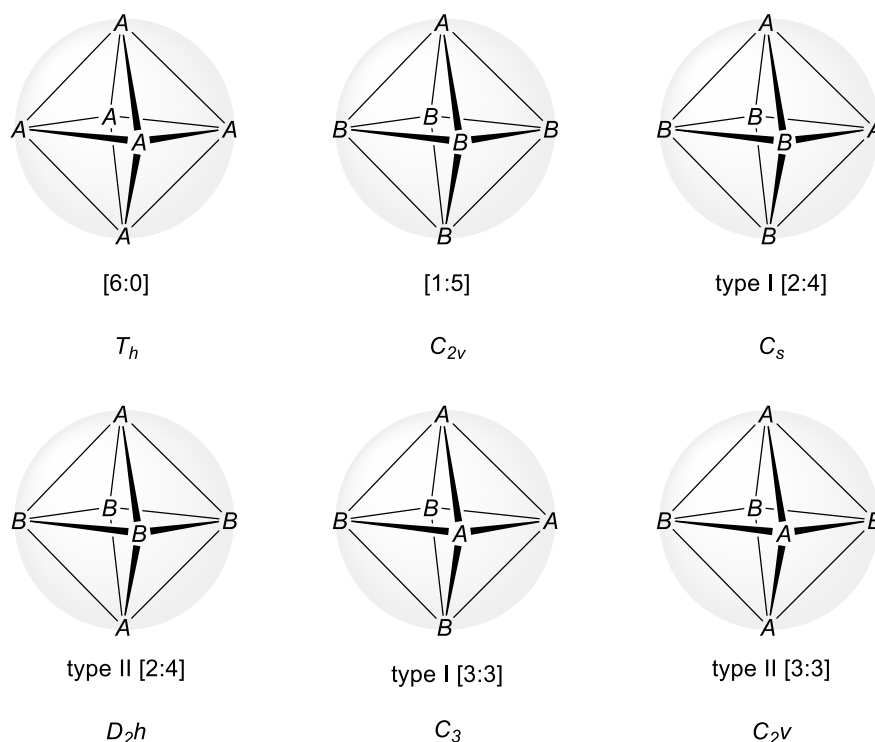


Figure IV-1 : Representation of the  $T_h$ -symmetrical hexa-adducts of  $C_{60}$ .

Hexa-adducts [6:0] and [1:5] are the easiest to prepare (Figure IV-2). The number in the bracket correspond to the different addends bond on  $C_{60}$  (e.g. [1:5] means that five malonates are identical and one malonate is different). Hexa-adducts [6:0] are directly prepared from  $C_{60}$  and hexa-adducts [1:5] from easily available  $C_{60}$  mono-adducts. In contrast, the preparation of more elaborated mixed hexa-adducts with different ratios of addends is more difficult. The preparation of fullerene bis- or tris-adduct precursors with a controlled regiochemistry is the key to have access to hexa-adducts with different ratio (e.g. [2:4] and [3:3])(Figure IV-2). Type I mixed hexa-adducts are obtained from lower multi-adducts with only *e* relationships. Type II mixed hexa-adducts are obtained from lower multi-adducts with *trans-1* and *e* relationships.<sup>[1]</sup>



*Figure IV-2 : Schematic representation of some mixed hexa-adducts of C<sub>60</sub>.*

Different strategies have been reported in the literature for the elaboration of mixed hexa-adducts and can be sorted in five different approaches.

*The stepwise approach:*

This approach is based on the reactivity of the *e* double bonds and no regioselective method is employed. Precursors were synthesized by stepwise procedures and required fastidious purifications of the *e* regioisomers. Mixed hexa-adducts could be obtained from the classical modified Bingel conditions.<sup>[2]</sup> This strategy is limited to symmetrical addends and to type I mixed hexa-adducts. Several mixed hexa-adducts prepared by stepwise approach have been described in the literature<sup>[1,3,4]</sup>.

*The template approach:*

This second approach is based on the addition reversibility of the Diels-Alder reaction and the reactivity of *e* double bonds. The regiochemistry is induced by the removable Diels-Alder adducts bond on the C<sub>60</sub>.

Kräutler and coworkers have developed a method to prepare a *trans-1* bis-adducts with removable anthracene addends.<sup>[5]</sup> This *trans-1* bisanthracene adduct can be used for the formation of type II [4:2] hexa-adducts with excellent yields (Figure IV-3). The two anthracene adducts were removed by heating the hexa-adducts in the solid state. The resulting tetra-adducts with an equatorial belt of addends could be then re-functionalized to afford type II fullerene [4:2] hexa-adducts.<sup>[6]</sup>

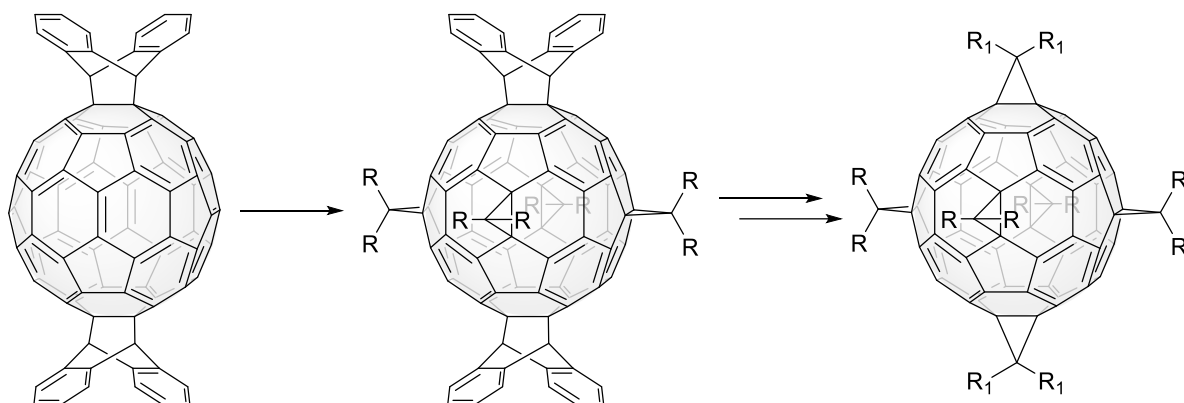


Figure IV-3 : Template strategy for the formation of type II [4:2] hexa-adducts.

The macrocyclic approach:

By a macrocyclic approach, multi-adducts of  $C_{60}$  can be formed with high regioselectivity. Using this strategy, only type I [3:3] hexa-adducts are reported in the literature.<sup>[7-9]</sup> The tris-adducts bearing a cyclo-[3]-octyl malonate have been used for the preparation of type I [3:3] hexa-adducts (Figure IV-4). The disadvantage of those compounds is that the cyclo-[3]-octyl malonate part cannot be subjected to further post-functionalization. This aspect limits considerably the scope of the functionalization of the carbon sphere.

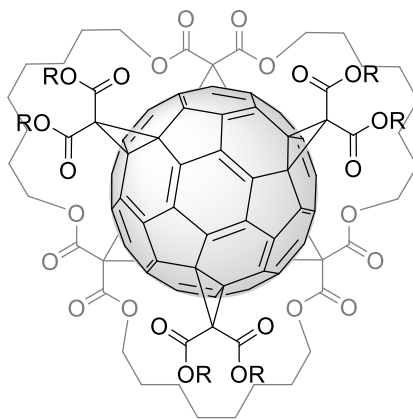


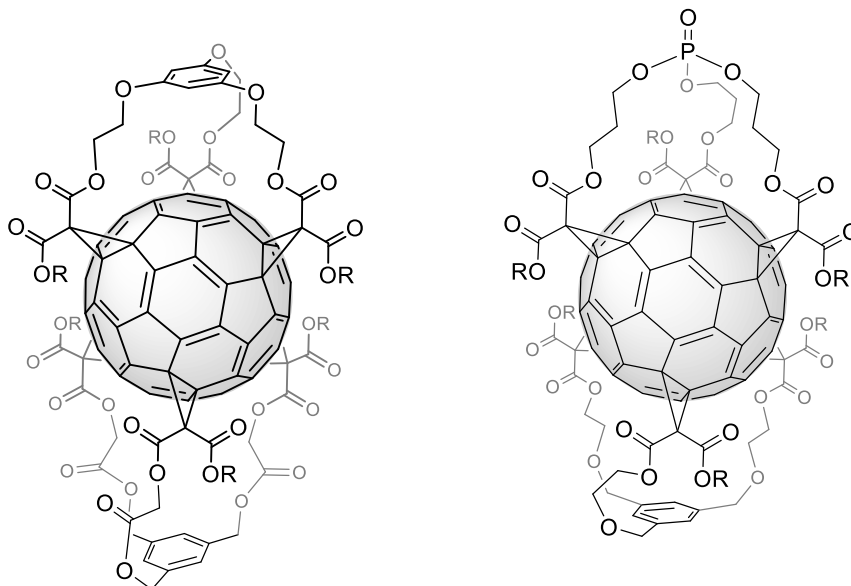
Figure IV-4 : Example of mixed  $C_{60}$  hexa-adduct obtained by the macrocyclic approach.

The tether-directed approach:

Currently, this is certainly the most efficient method to control the regio- and the stereo-chemistry of the addends around the  $C_{60}$ . The design of the tether is an important matter. Indeed, removable of the tether would allow post-functionalization of the addends and thus afford building blocks for the preparation of new compounds with specific properties.

Hirsch and coworkers have developed three different tripodal malonate tethers for the regioselective preparation of tris-adducts of  $C_{60}$  in order to have access to mixed [3:3] hexa-

adducts of  $C_{60}$  with four spherically defined addend zones ([Figure IV-5](#)).<sup>[10,11]</sup> The removal of two of the three tethers afforded carboxylic acid, alcohol or bromide functions depending on the used tether. Different reactions can be envisaged based on the reactivity of the released functions for the post-functionalization of the hexa-adducts.



[Figure IV-5](#) : The three tethers developed by Hirsch and coworkers for the preparation of mixed [3:3] hexa-adducts of  $C_{60}$ .

#### Mixed approach:

This category regroups the mixed hexa-adducts formed by the union of the tether-directed, template and stepwise approaches.

Diederich and coworkers have developed a malonate tether bearing two “Diels-Alder addends”. Reaction of this tether with  $C_{60}$  gave only one regioisomer of tris-adducts with a  $C_{2v}$  symmetry which was assigned to an *e,e,trans-1* tris-adducts of  $C_{60}$ . The two Diels-Alder adducts have been removed by retro-Diels-Alder reactions after the formation of the  $C_{60}$  hexa-adducts ([Figure IV-6](#)). Following different paths, they were able to synthesize mixed hexa-adducts with different ratio bearing up to four different addends (e.g. type II [4:2], [1:2:3] and [1:1:2:2]).<sup>[12–16]</sup>

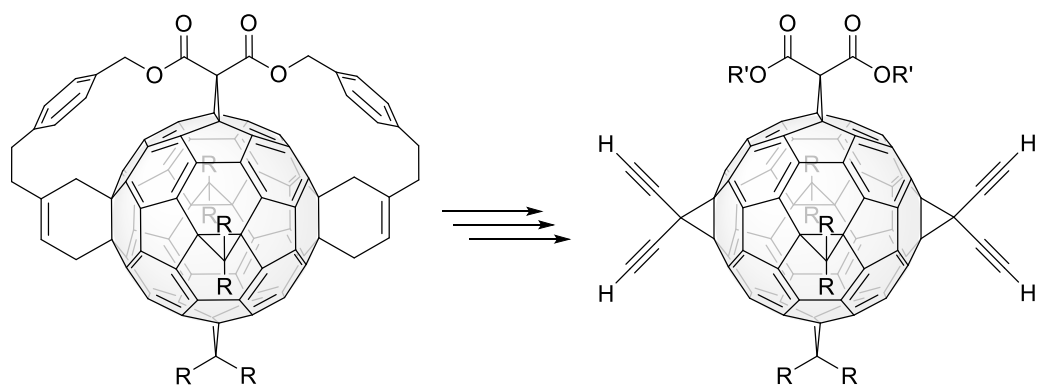


Figure IV-6 : Strategy employed by Diederich and coworkers for the preparation of mixed hexa-adducts of C<sub>60</sub>.

Rubin and coworkers used a tether with two “Diels-Alder addends” at both ends. They obtained a *trans*-1 bis-adducts with a masked *e* position by the tether. The complete functionalization of the carbon sphere led only to the penta-adducts with an incomplete octahedral addition pattern (Figure IV-7). Mixed [3:3] hexa-adducts were obtained by the removal of the tether and the subsequent three-fold additions. It was also observed that the remaining *e* positions could be exclusively functionalized in a sequential manner (*e*-face, *e*-edge and finally *e'*-face).<sup>[17]</sup> In the continuity of this work, they succeeded to synthesize seven mixed [2:2:2] C<sub>60</sub> hexa-adducts out of the eight possible regioisomers.<sup>[18]</sup>

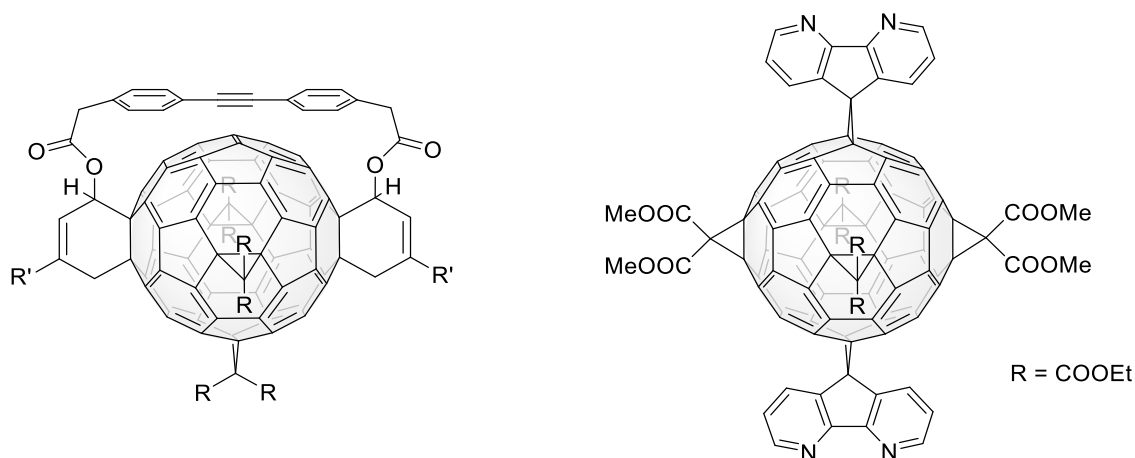


Figure IV-7 : left: Penta-adduct with an incomplete octahedral addition pattern; right: Example of one of the seven mixed [2:2:2] hexa-adducts obtained.

In the reported methodologies, the lack of post-functionalization possibilities, low yields or fastidious syntheses limited the preparation of highly functionalized mixed hexa-adducts of C<sub>60</sub>. In this chapter, we will exploit the synthetic strategies described in the previous chapters in order to synthesize mixed C<sub>60</sub> hexa-adducts. These compounds will be furthermore post-functionalized by sequential “click” reactions in order to obtain multi-functionalized C<sub>60</sub> hexa-adducts.

## 2. Synthesis of mixed C<sub>60</sub> hexa-adducts by “click” reactions.

### 2.1 Preparation of mixed C<sub>60</sub> hexa-adducts building blocks.

Based on our efficient and regioselective methodology for the preparation of *e,e,e* tris-adducts, we have now easy access to the triol building block **III-14** in a relatively good yield (13% over 4 steps). With its *e,e,e* addition pattern, compound **III-14** is perfectly suited for the preparation of mixed hexa-adducts with an octahedral addition pattern. In order to have access to more functionalized hexa-adducts, the ethyl esters of **III-14** must be modified to allow the post-functionalization of these side chains. For this reason, malonate **IV-2** was prepared from 3-bromo-1-propanol (Scheme IV-1) in order to obtain two more building blocks: the *e,e,e* C<sub>60</sub> tris-adduct **IV-6** and the *e* C<sub>60</sub> bis-adduct **IV-9** (Figure IV-8).

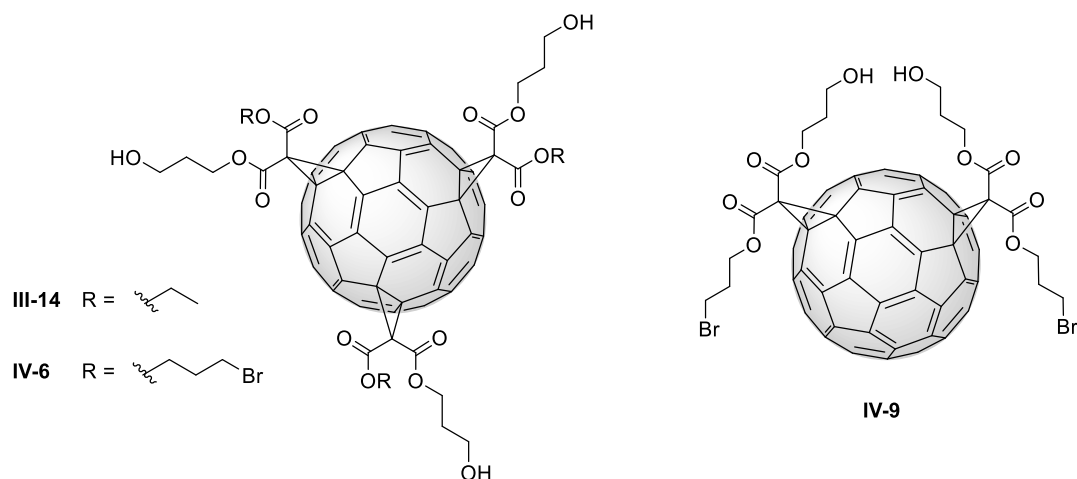
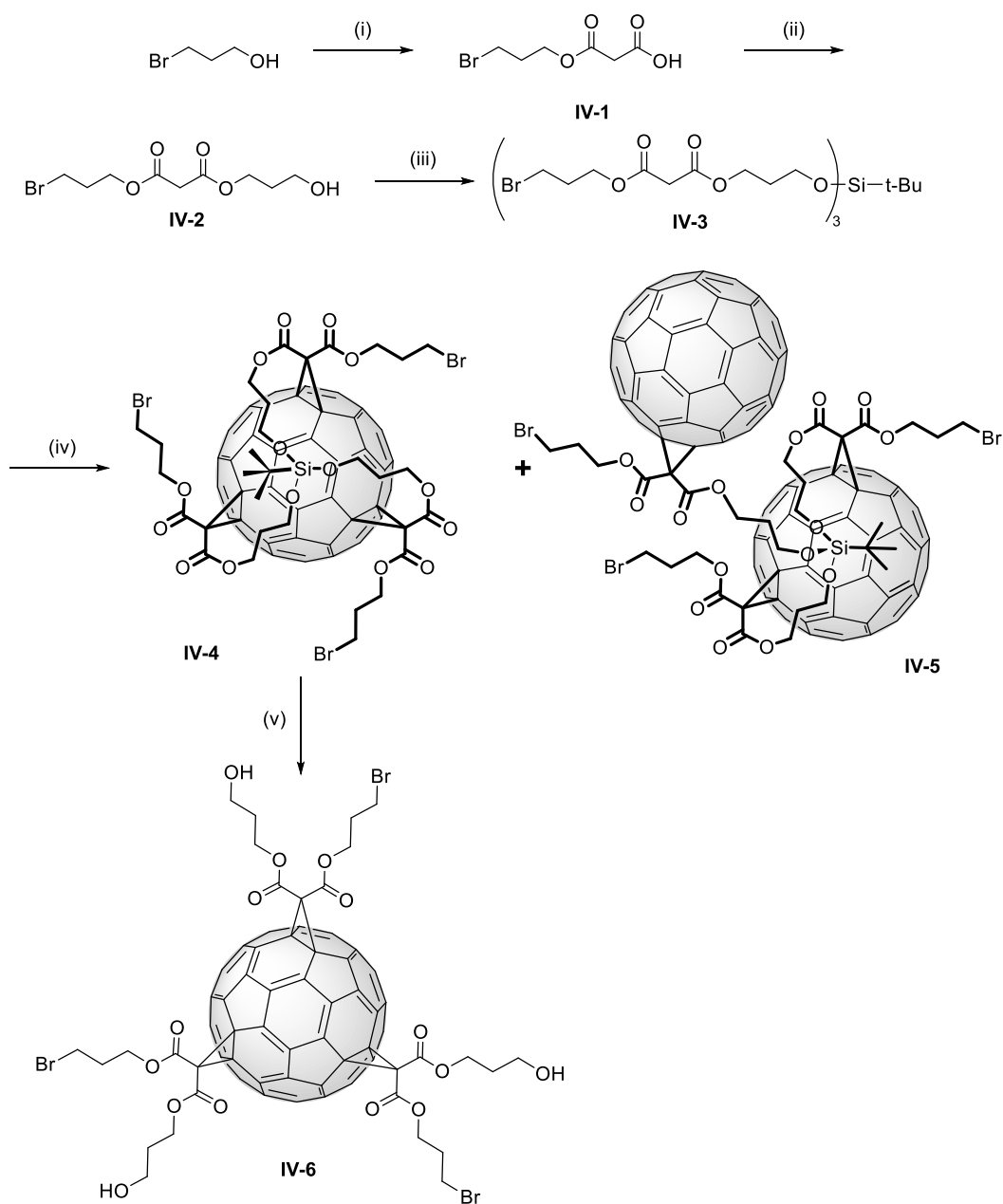


Figure IV-8 : The three building blocks used for the preparation of mixed C<sub>60</sub> hexa-adducts.

Reaction of 3-bromo-1-propanol with Meldrum acid at 80 °C afforded the malonic acid derivative **IV-1** (Scheme IV-1). Subsequent esterification reaction of **IV-1** with an excess of 1,3-propanediol in the presence of DCC and DMAP gave the malonate **IV-2** in 46% yield over two steps. The tris-malonate derivative **IV-3** was then prepared by treatment of **IV-2** with *t*-BuSiCl<sub>3</sub> in the presence of pyridine in CH<sub>2</sub>Cl<sub>2</sub> at 0 °C. The key conditions of this reaction were the use of an apolar solvent and a low temperature to avoid bromide to chloride substitution side reactions. Reaction of **IV-3** with C<sub>60</sub>, under the conditions developed for preparation of fullerene *e,e,e* tris-adducts from *t*-butyl(trialkoxy)silane derivatives, afforded tris-adduct **IV-4** and bis-fullerene derivative **IV-5**. The side chains of the tris-malonate **IV-3** do not affect the regioselectivity of the reaction and the *e,e,e* addition pattern was confirmed by NMR and UV/vis spectroscopies. The structure of **IV-4** was further confirmed by mass spectrometry. Finally, treatment of **IV-4** with BF<sub>3</sub>.Et<sub>2</sub>O (20 equiv) in CH<sub>2</sub>Cl<sub>2</sub>/CH<sub>3</sub>CN afforded the triol derivative **IV-6** in a good yield. The protons of the alkyl chains are diastereotopic as observed on the <sup>1</sup>H



NMR spectrum of **IV-6**. That can be rationalized by intramolecular hydrogen-bonding between the released alcohol functions (**Figure IV-9**).<sup>[19]</sup>



**Scheme IV-1.** Reagents and conditions: (i) Meldrum acid, 80 °C; (ii) 1,3-propanediol, DCC, DMAP,  $\text{CH}_2\text{Cl}_2$ , 0 °C to rt, 12 h (46% over two steps); (iii)  $t\text{BuSiCl}_3$ , pyridine,  $\text{CH}_2\text{Cl}_2$ , rt, 12 h (66%); (iv)  $\text{C}_{60}$ ,  $\text{I}_2$ , DBU, PhMe, -15 °C, 1 h (**IV-4**: 13% and **IV-5**: 3%); (v)  $\text{BF}_3 \cdot \text{Et}_2\text{O}$ ,  $\text{CH}_2\text{Cl}_2/\text{MeCN}$  2:1, rt, 12 h (86%).

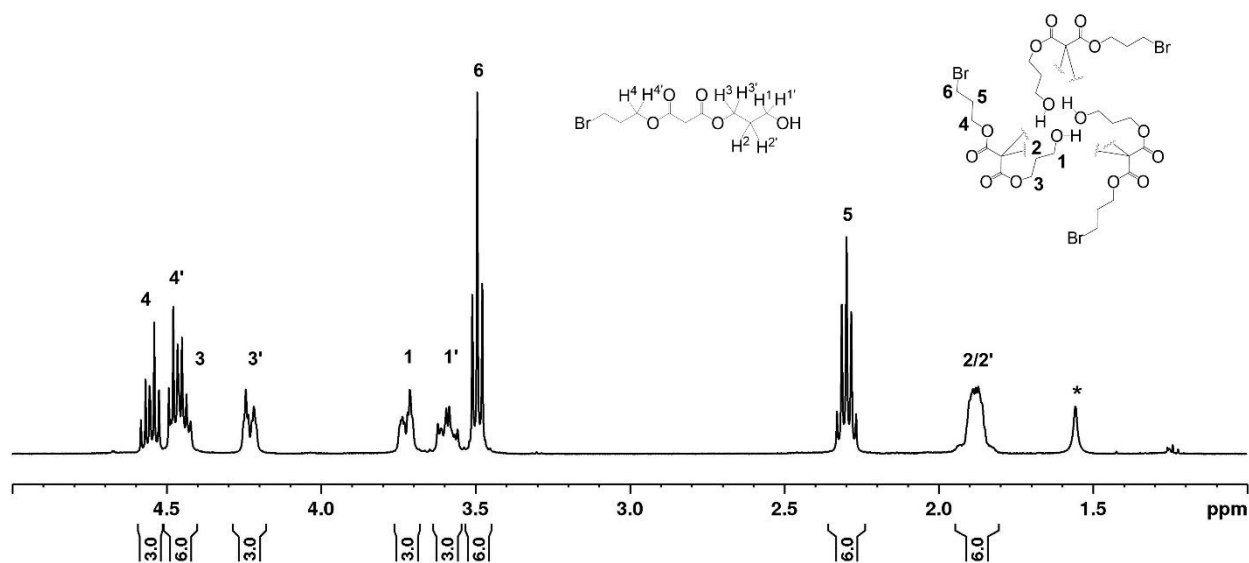
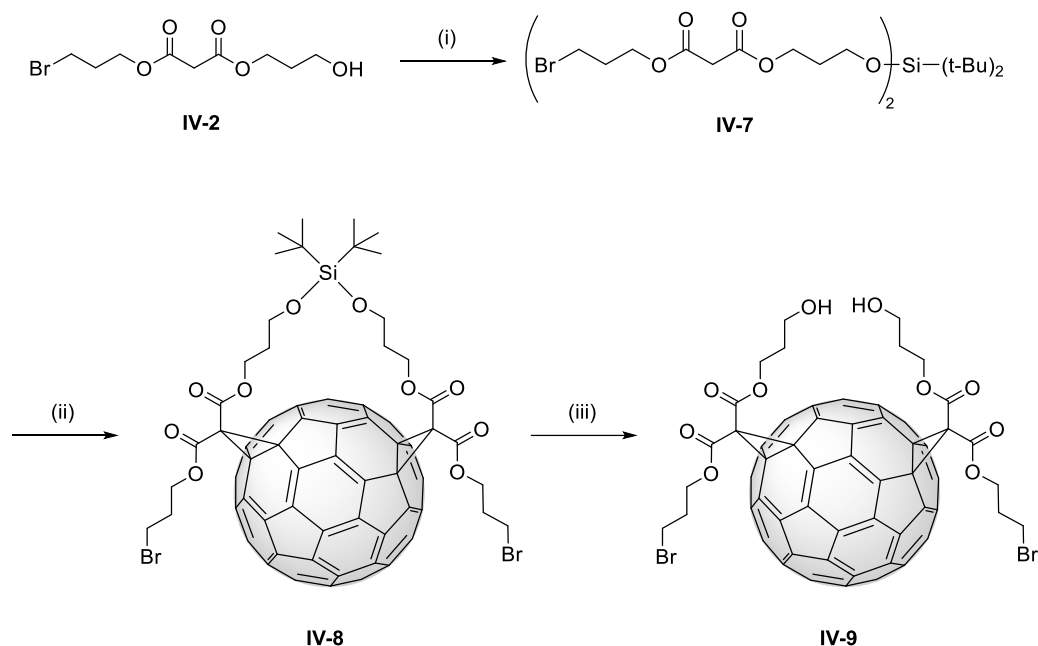


Figure IV-9 :  $^1\text{H}$  NMR spectrum (400 MHz,  $\text{CDCl}_3$ ) of **IV-6**. (\*) water.

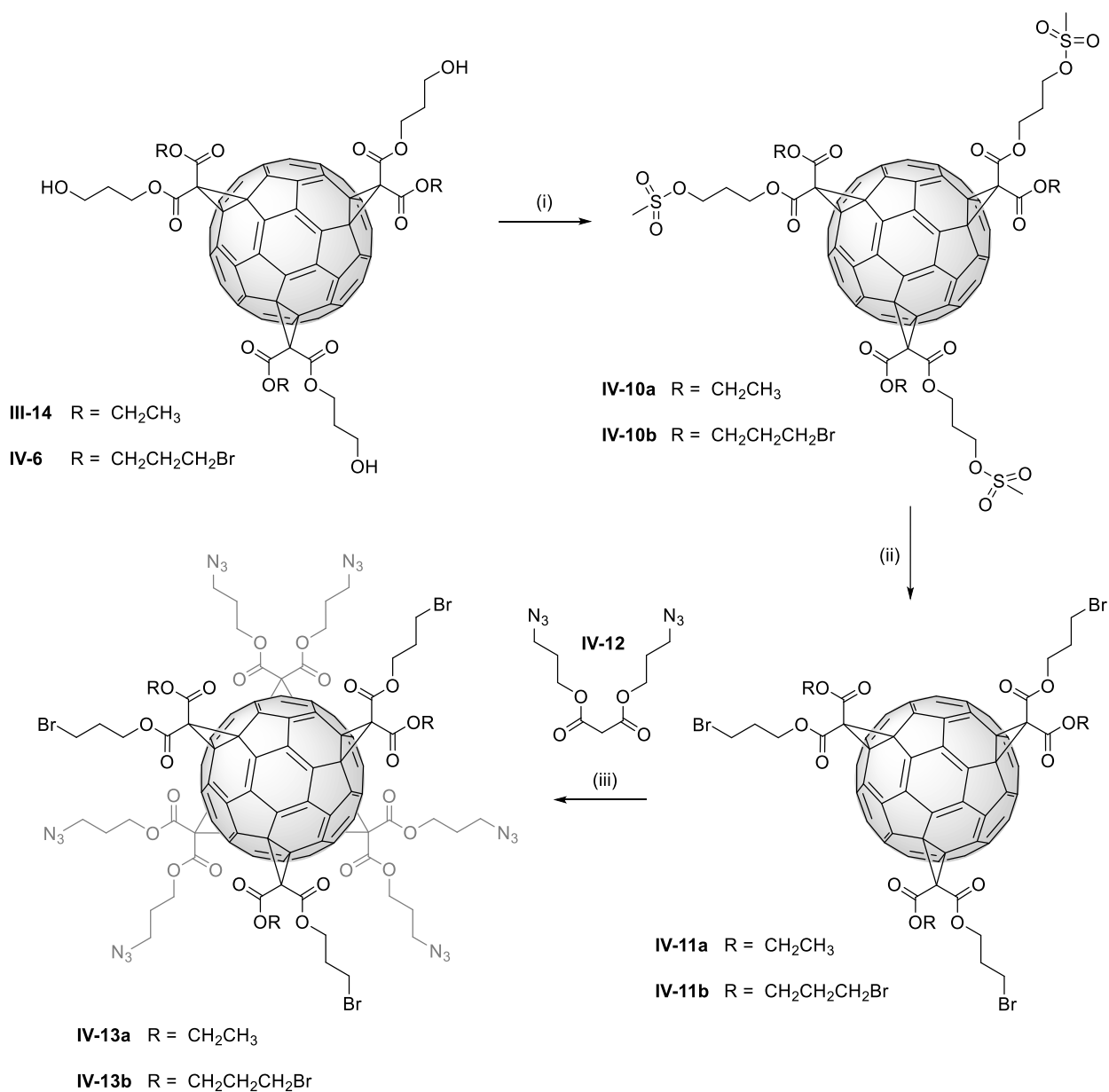
The synthesis of compound **IV-9** is depicted in **Scheme IV-2**. Treatment of compound **IV-2** (2 equiv.) with di-*t*-butylsilyl bis(trifluoromethanesulfonate) ( $t\text{Bu}_2\text{Si}(\text{OTf})_2$ , 1 equiv.) in DMF in the presence of pyridine gave the bis-malonate **IV-7**. Reaction of **IV-7** with  $\text{C}_{60}$ ,  $\text{I}_2$  and DBU in toluene afforded the corresponding cyclization product **IV-8** in 48% yield. Fullerene bis-adduct **IV-8** was characterized by  $^1\text{H}$  and  $^{13}\text{C}$  NMR, UV/vis and IR spectroscopies. In addition, its structure was confirmed by MALDI-TOF mass spectrometry showing the expected pseudo-molecular ion peaks at  $m/z = 1423.0$  ( $[\text{M}+\text{H}]^+$ , calcd for  $\text{C}_{86}\text{H}_{43}\text{O}_{10}\text{Br}_2\text{Si}$ : 1423.1). The  $\text{C}_1$  molecular symmetry deduced from the  $^1\text{H}$  and  $^{13}\text{C}$  NMR spectra suggests an *equatorial* addition pattern. This was further confirmed by the UV/vis spectrum revealing the diagnostic features previously reported for analogous *equatorial*  $\text{C}_{60}$  bis-adducts. Finally, desilylation of **IV-8** with  $\text{BF}_3 \cdot \text{Et}_2\text{O}$  provided *e* bis-adduct **IV-9**.



**Scheme IV-2.** Reagents and conditions: (i)  $(t\text{Bu})_2\text{Si}(\text{OTf})_2$ , pyridine, DMF, rt, 12 h (68%); (ii)  $\text{C}_{60}$ ,  $\text{I}_2$ , DBU, PhMe, rt, 1 h (48%); (iii)  $\text{BF}_3 \cdot \text{Et}_2\text{O}$ ,  $\text{CH}_2\text{Cl}_2/\text{MeCN}$  2:1, rt, 12 h (91%).

The post-functionalization of  $\text{C}_{60}$  hexa-adducts by copper-catalyzed alkyne-azide 1,3-dipolar cycloaddition (CuAAC) reactions has shown its efficiency<sup>[20]</sup> and thus, we oriented the synthesis of the mixed hexa-adducts building blocks in this direction.

As shown in **Scheme IV-3**, the preparation of tris-adduct **IV-11** was first achieved. Treatment of **III-14** and **IV-6** with methanesulfonyl chloride ( $\text{MeSO}_2\text{Cl}$ ) in  $\text{CH}_2\text{Cl}_2$  in the presence of triethylamine at  $0^\circ\text{C}$  gave compounds **IV-10a,b**. As for the preparation of compound **IV-3**, the low temperature was the key condition to avoid the substitution of the bromide by a chloride. Subsequent bromination of **IV-10** with LiBr in THF afforded **IV-11** in good yields. Reaction of **IV-11a-b** with **IV-12** (5 equiv.) were then performed in the presence of  $\text{CBr}_4$  (100 equiv.) and DBU (18 equiv.) in toluene at room temperature.<sup>[2]</sup> The  $\text{C}_3$ -molecular symmetry deduced from the  $^1\text{H}$  NMR spectrum ([Figure IV-10](#)) is in agreement with the octahedral addition pattern of  $\text{C}_{60}$  hexa-adducts **IV-13**. In the case of **IV-13**, the reactivity of azides towards the  $\text{C}_{60}$  core is considerably reduced but reactions can still occur. <sup>[11,21,22]</sup> Typically,  $\text{C}_{60}$  hexa-adducts bearing azides are rapidly used after purification.



**Scheme IV-3.** Reagents and conditions: (i) MeSO<sub>2</sub>Cl, Et<sub>3</sub>N, CH<sub>2</sub>Cl<sub>2</sub>, 0 °C, 2 h (**IV-10a**: 86%; **IV-10b**: 94%); (ii) LiBr, THF, 60 °C, 12 h (**IV-11a**: 83%; **IV-11b**: 98%); (iii) **IV-12**, CBr<sub>4</sub>, DBU, PhMe, rt, 12 h (**IV-13a**: 42%; **IV-13b**: 63%).

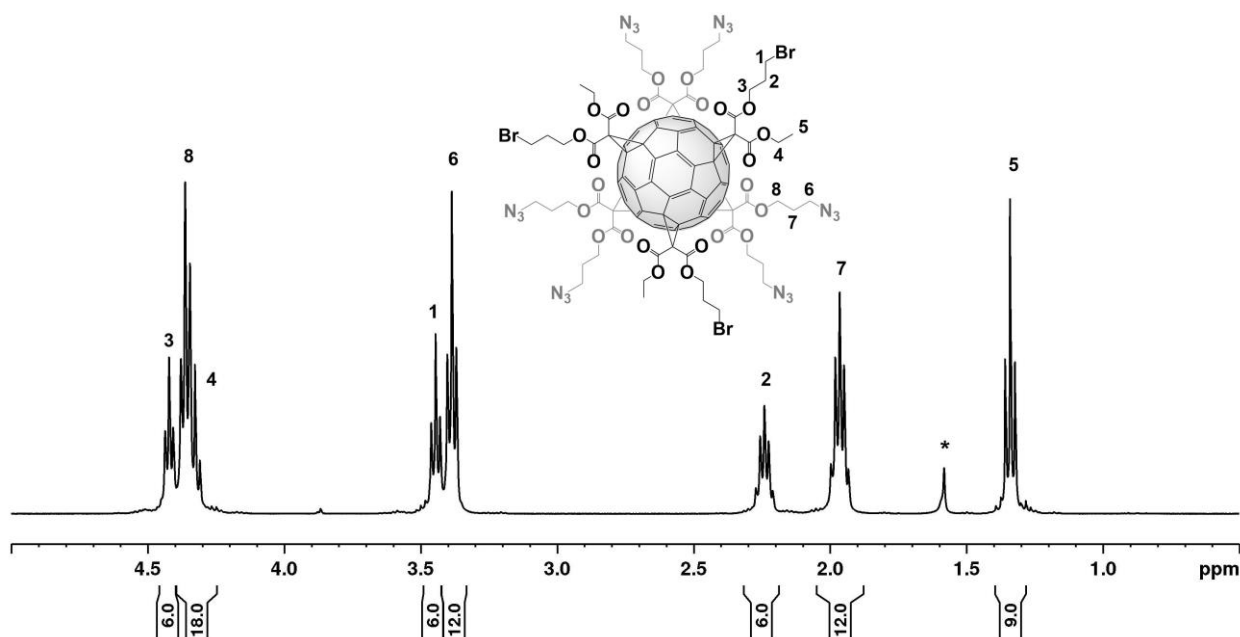
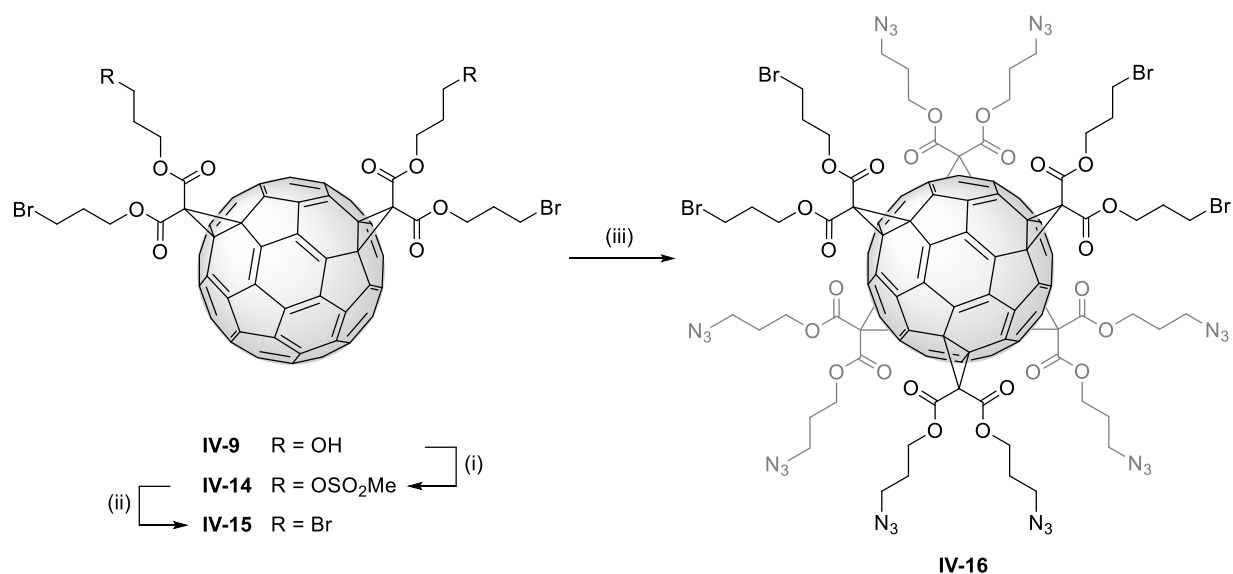
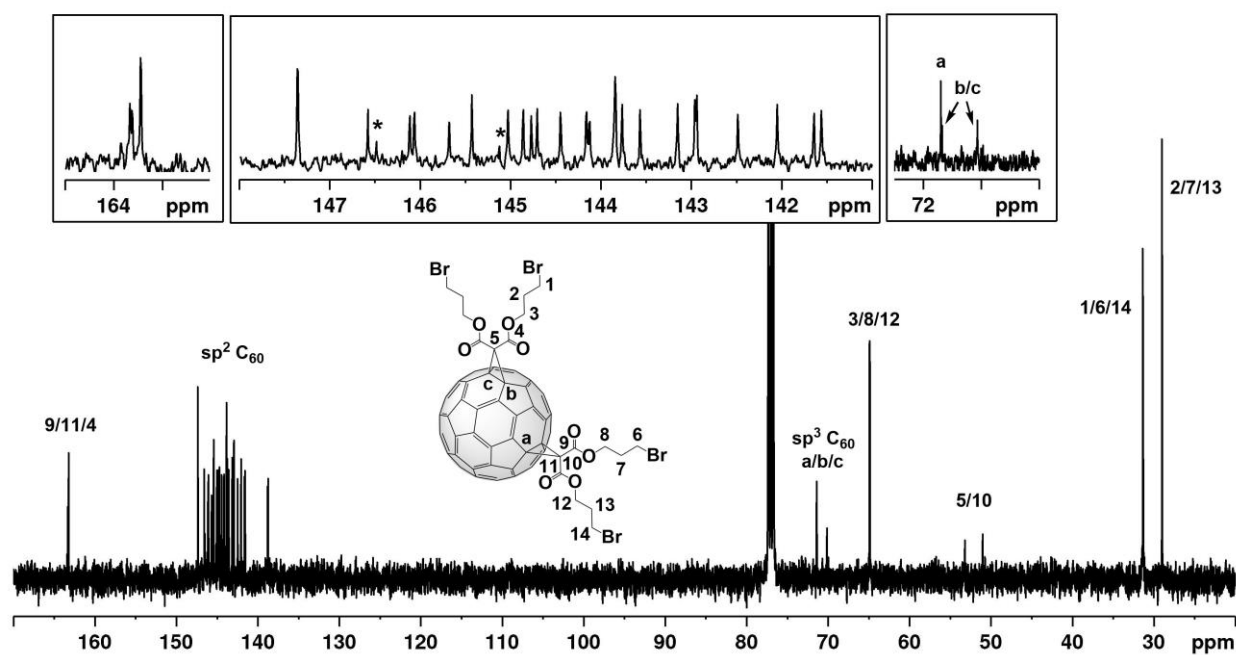


Figure IV-10:  $^1\text{H}$  NMR spectrum (400 MHz,  $\text{CDCl}_3$ ) of **IV-13a** showing the  $C_3$  molecular symmetry. (\*) water.

Following the same synthetic route, the preparation of the mixed [2:4]  $C_{60}$  hexa-adduct **IV-16** was achieved (**Scheme IV-4**). Mesylation of **IV-9** followed by bromination afforded the tetrabrominated bis-adduct **IV-15**. Whereas derivatives **IV-8**, **IV-9** and **IV-14** bearing unsymmetrically substituted malonate addends are all  $C_1$ -symmetrical compounds,  $C_{60}$  bis-adduct **IV-15** with its two identical malonate subunits is  $C_5$  symmetric as deduced from a careful analysis of its  $^{13}\text{C}$  NMR spectrum (**Figure IV-11**). Out of the 32 expected fullerene resonances, three are observed at  $\delta = 70.1$ ,  $71.3$  and  $71.4$  ppm ( $\text{sp}^3$  C atoms) and 29 between  $\delta = 138.7$  and  $147.4$  ppm ( $\text{sp}^2$  C atoms). It is also worth noting that two of the resonances seen in the  $\text{sp}^2$  region show half intensity as well as two of the three resonances observed for the fullerene  $\text{sp}^3$  C atoms. Actually, the latter observation are unambiguous proofs for the  $C_5$  symmetry of compound **IV-15**. Furthermore, out of the three possible  $C_5$ -symmetrical addition patterns (cis-2, e and trans-4), the only one for which 3 resonances are expected for the  $\text{sp}^3$  fullerene C atoms is the *equatorial* bis-adduct. This is further confirmed by the observation of three resonances for the carbonyl C atoms ( $\delta = 163.2$ ,  $163.29$  and  $163.32$  ppm). Finally, formation of the hexa-adduct **IV-16** was achieved in 56% yield.



**Scheme IV-4.** Reagents and conditions: (i) MeSO<sub>2</sub>Cl, Et<sub>3</sub>N, CH<sub>2</sub>Cl<sub>2</sub>, 0 °C, 15 min (93%); (ii) LiBr, THF, 60 °C, 12 h (96%); (iii) **IV-12**, CBr<sub>4</sub>, DBU, PhMe, rt, 12 h (56%).



**Figure IV-11 :**  $^{13}\text{C}$  NMR spectrum (100 MHz, CDCl<sub>3</sub>) of **IV-15** showing the  $\text{C}_s$ -symmetry; Inset: detailed views showing the resonances of the carbonyl and the fullerene  $\text{sp}^2$  and  $\text{sp}^3$  C atoms; \* indicates the  $\text{sp}^2$  fullerene C atoms showing half intensity signals.

## 2.2 Post-functionalization of the mixed C<sub>60</sub> hexa-adducts by CuAAC reactions.

Based on the three developed hexa-adducts building blocks **IV-13a,b** and **IV-16**, two different addend zones on the C<sub>60</sub> can be functionalized by sequential CuAAC reactions.

The synthesis of the mixed [3:3] C<sub>60</sub> hexa-adduct **IV-19** was performed first (**Scheme IV-5**). Reaction of **IV-13a** with phenylacetylene under typical conditions of CuAAC reactions (CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate and CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O) gave the first “click” compound **IV-17** in 94% yield (**Scheme IV-5**). Compound **IV-17** has been characterized by <sup>1</sup>H and <sup>13</sup>C NMR, UV/vis and IR spectroscopies. The absence of azide residues was confirmed by the IR data (2099 cm<sup>-1</sup>) and by the disappearance of the CH<sub>2</sub>-N<sub>3</sub> signal at δ = 3.39 ppm on the <sup>1</sup>H NMR spectrum (**Figure IV-12**). The <sup>1</sup>H NMR spectrum of **IV-17** confirmed also the formation of the triazoles with the presence of the characteristic singlets of the 1,2,3-triazole units at δ = 7.82 and 7.86 ppm as well as the signal of the CH<sub>2</sub>-triazole protons at δ = 4.4 ppm (**Figure IV-12**). In addition, the MALDI-TOF mass spectrum shows the expected pseudo-molecular ion peak at *m/z* = 2892.8 ([M+H]<sup>+</sup>, calcd for C<sub>159</sub>H<sub>106</sub>O<sub>24</sub>N<sub>18</sub>Br<sub>3</sub>: 2892.5).

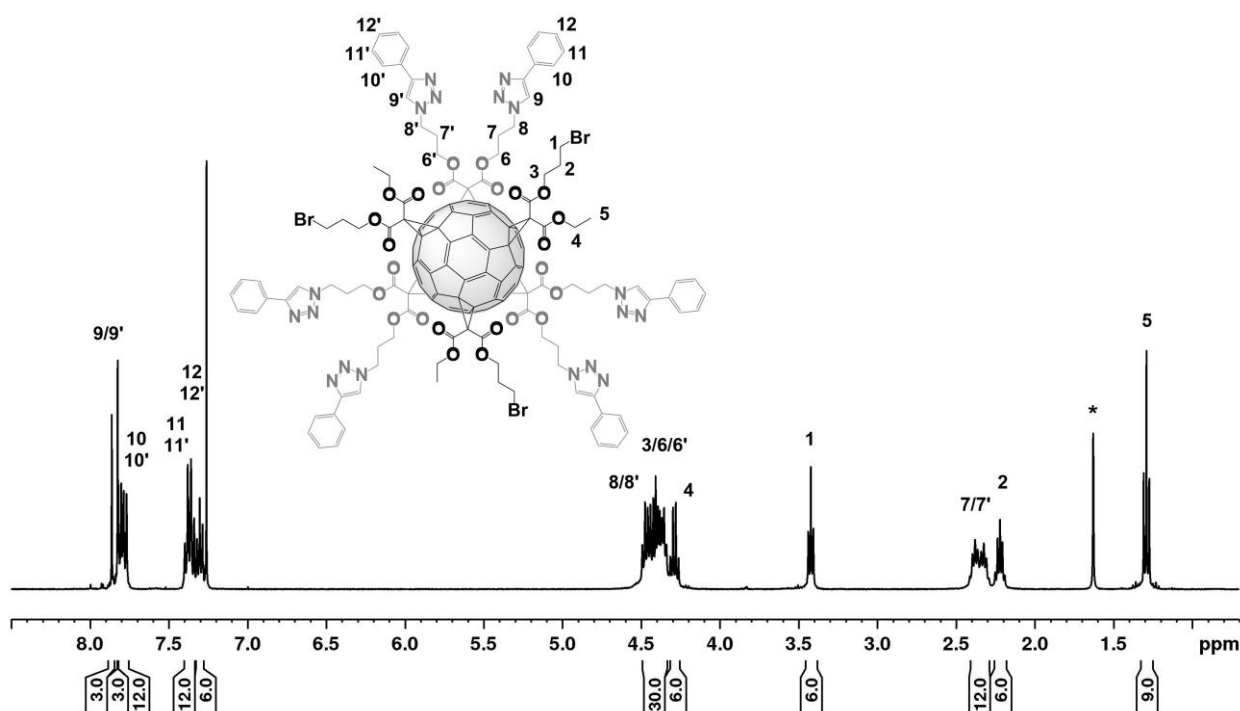
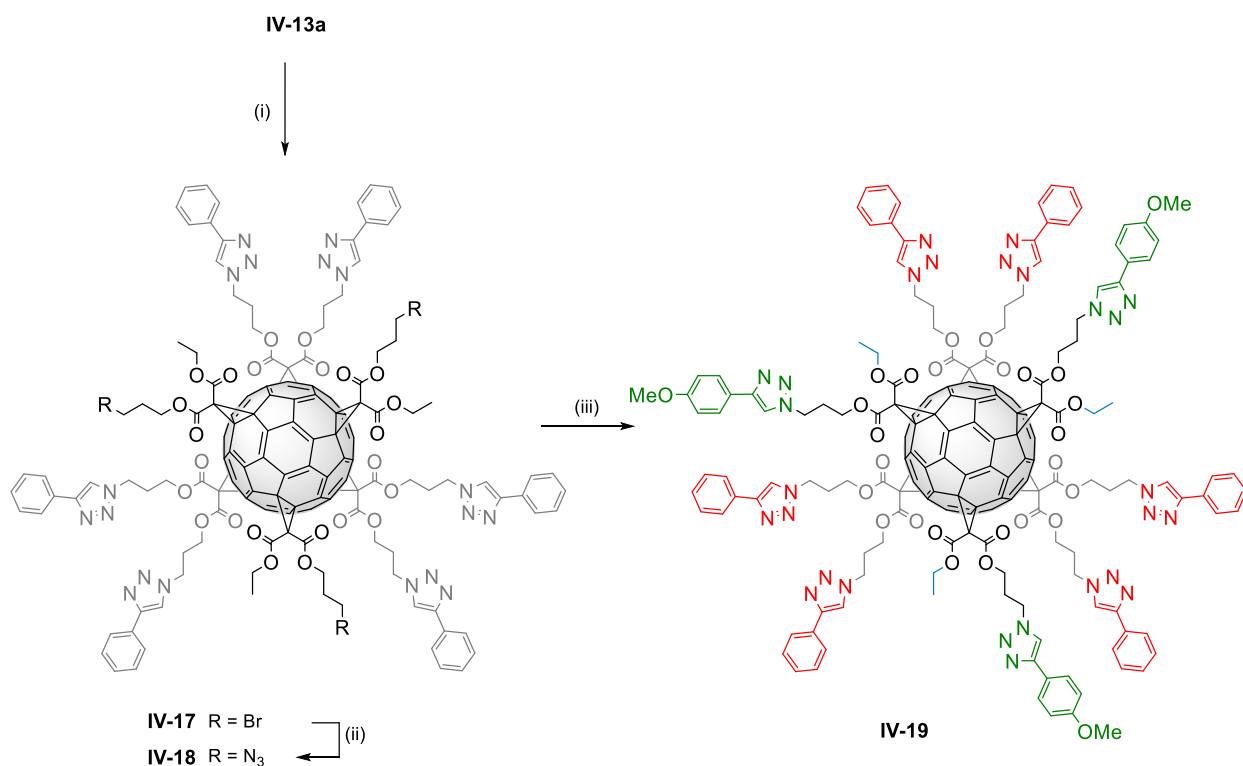


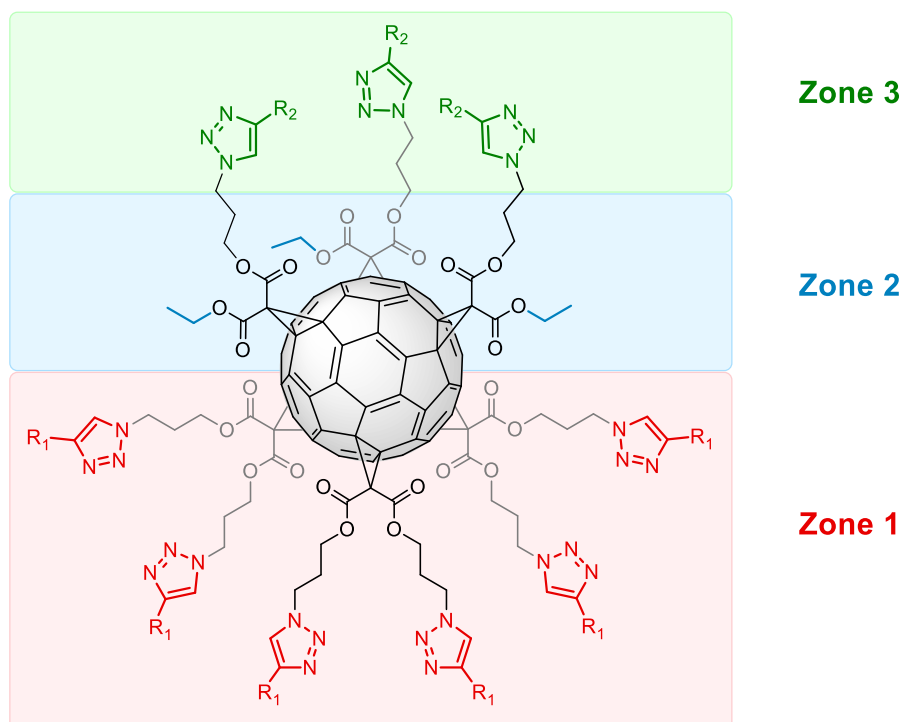
Figure IV-12 : <sup>1</sup>H NMR spectrum (400 MHz, CDCl<sub>3</sub>) of **IV-17**. (\*) water.



**Scheme IV-5.** Reagents and conditions: (i) Phenylacetylene, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (94%); (ii) NaN<sub>3</sub>, DMF, rt, 12 h (96%); (iii) 4-ethynylanisole, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 19 h (97%).

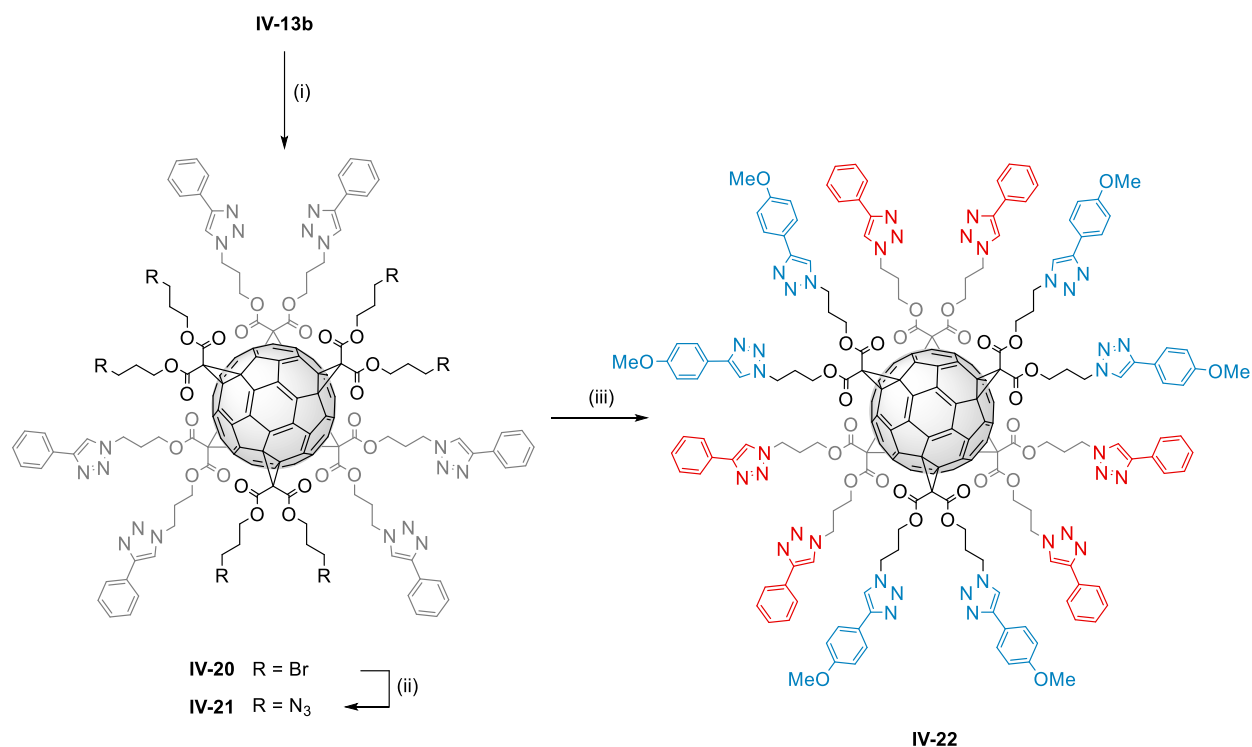
Subsequent conversion of the bromide functions in **IV-17** by treatment with sodium azide in DMF afforded tris-azide **IV-18**. Finally, CuAAC reaction of **IV-18** with 4-ethynylanisole gave mixed hexa-adduct **IV-19** in an excellent yield (97%). The chemical structure of **IV-19** was confirmed by <sup>1</sup>H and <sup>13</sup>C NMR, UV/vis and IR spectroscopies, mass spectrometry and elemental analysis. The mixed [3:3] hexa-adducts **IV-19** with a function ratio of (6:3) was obtained in 10 steps from 1,3-propanediol in an overall yield of 3%. Three different addend zones can be defined in the mixed hexa-adducts **IV-19** by the control of the regio- and the stereochemistry of our methodology (Figure IV-13).



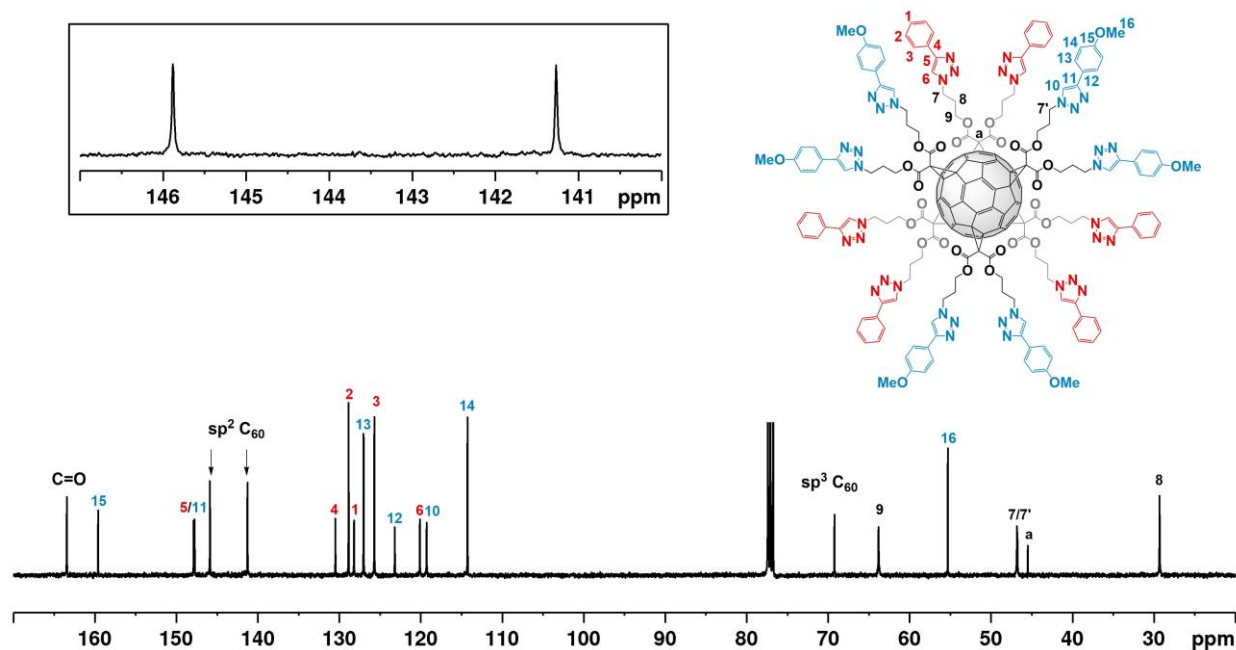


*Figure IV-13 : Illustration of the three different addend zones of the mixed hexa-adduct IV-19.*

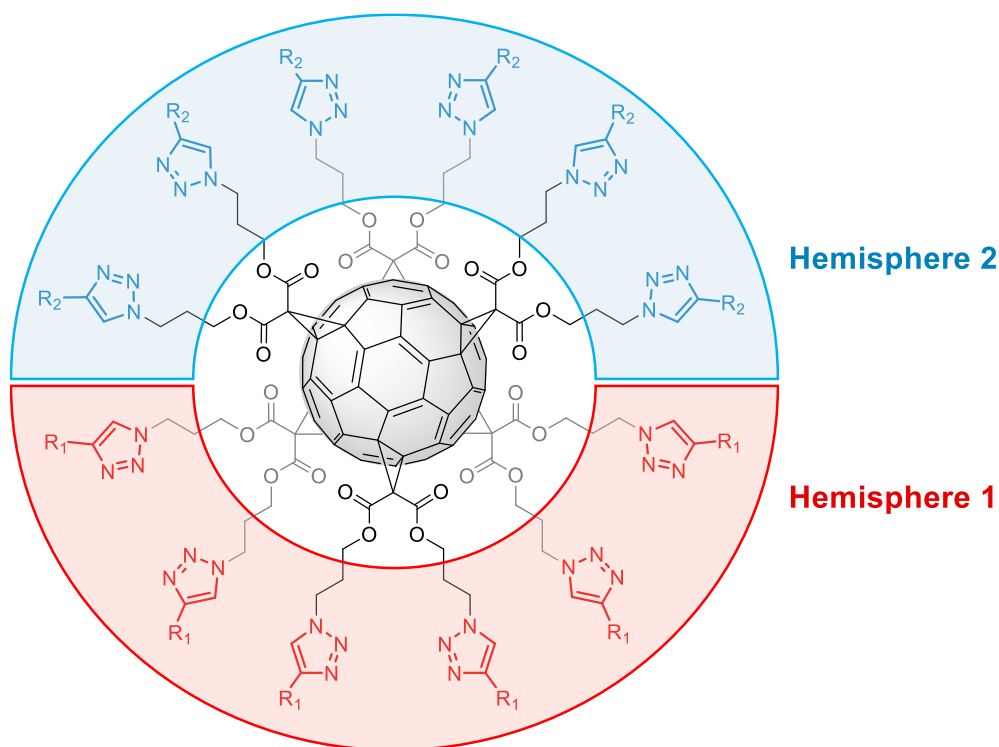
Following the successful preparation of hexa-adducts **IV-19** from the triol **III-14**, the same synthetic route was used for the preparation of an additional mixed [3:3] hexa-adduct **IV-22** with a (6:6) function ratio (**Scheme IV-6**). The final hexa-adduct **IV-22** was synthesized from compound **IV-13b** by two sequential CuAAC reactions. The chemical structure of **IV-22** was confirmed by  $^1\text{H}$  and  $^{13}\text{C}$  NMR, UV/vis and IR spectroscopies, mass spectrometry and elemental analysis. As shown in [Figure IV-14](#), the  $^{13}\text{C}$  NMR spectrum of **IV-22** exhibits a high local symmetry with only three fullerene resonances at  $\delta = 145.9$  and  $141.3$  ppm for the  $\text{sp}^2$  C atoms and at  $\delta = 69.2$  ppm for the  $\text{sp}^3$  C atoms which is in full agreement with an octahedral addition pattern. No influence of the  $\text{C}_3$  overall symmetry of **IV-22** can be deduced for the  $\text{C}_{60}$  core. Similar observations have been reported by Hirsch and coworkers for mixed hexa-adducts of  $\text{C}_{60}$ .<sup>[1]</sup> The particularity of this mixed [3:3] hexa-adduct **IV-22** is the localization of the two different functions in two defined hemispheres ([Figure IV-15](#)).



**Scheme IV-6.** Reagents and conditions: (i) Phenylacetylene, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (99%); (ii) NaN<sub>3</sub>, DMF, rt, 12 h (98%); (iii) 4-ethynylanisole, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (88%).



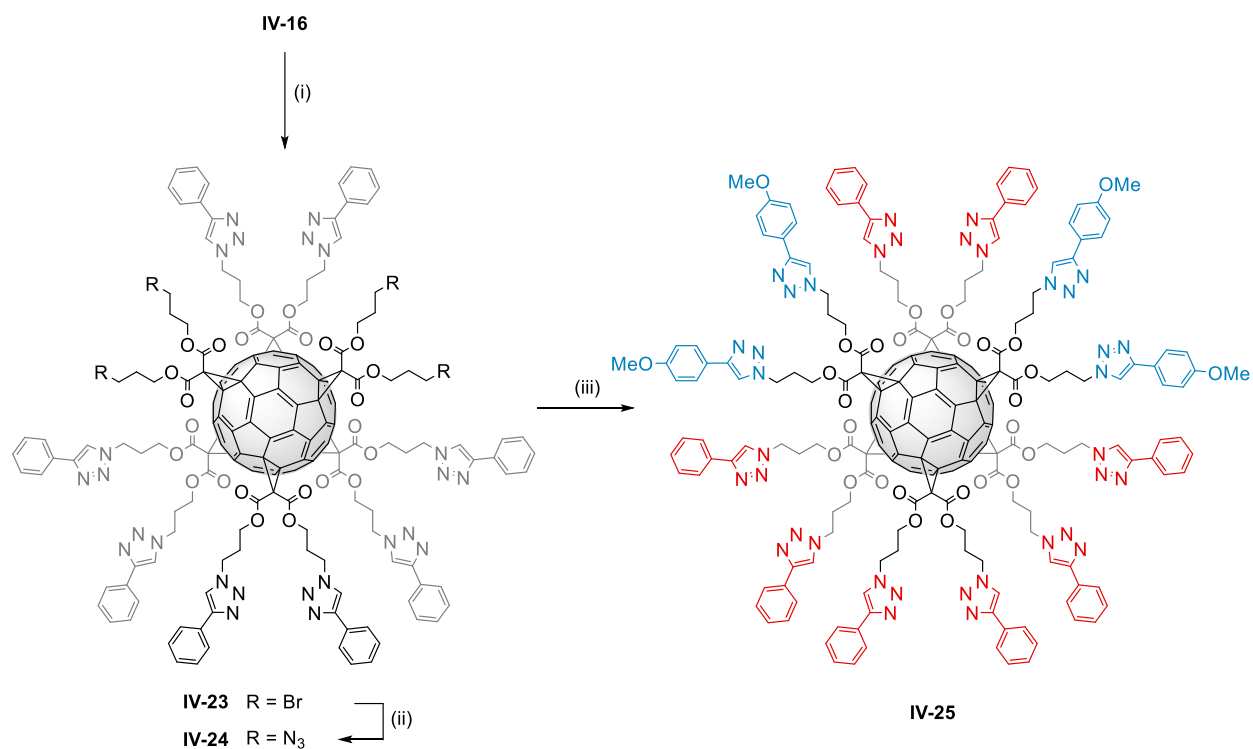
**Figure IV-14:** <sup>13</sup>C NMR spectrum (100 MHz, CDCl<sub>3</sub>) of IV-22 showing the high local symmetry. Inset: detailed view showing the resonances of the fullerene sp<sup>2</sup> C atoms.



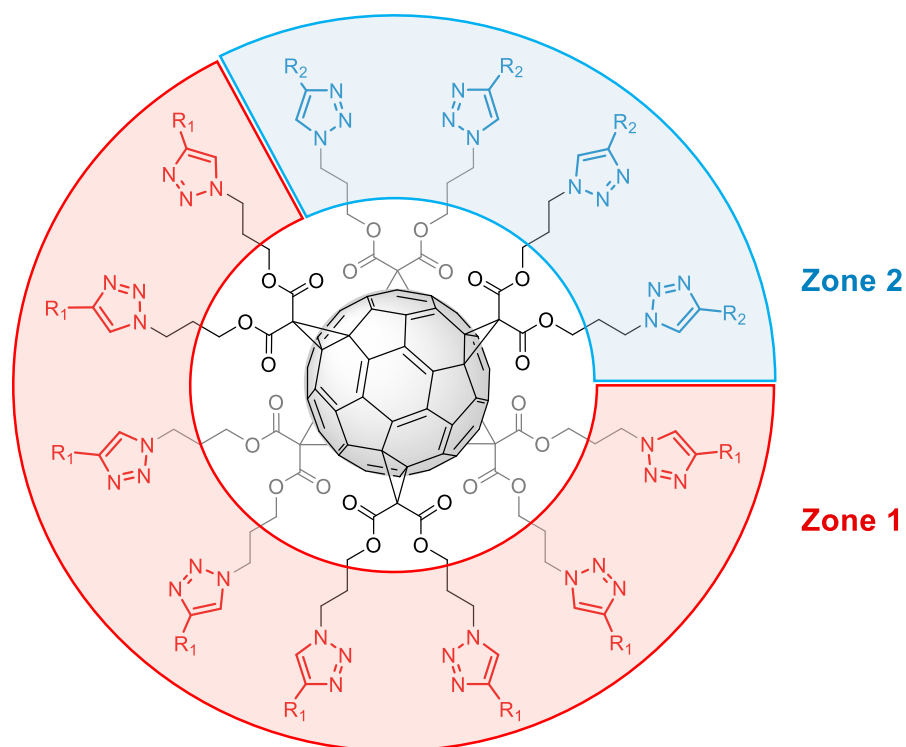
*Figure IV-15 : Illustration of the two hemisphere of the mixed C<sub>60</sub> hexa-adduct IV-22.*

Finally, the [2:4] hexa-adduct **IV-25** was also synthesized by two sequential CuAAC reactions (**Scheme IV-7**). The chemical structure of **IV-25** was confirmed by <sup>1</sup>H and <sup>13</sup>C NMR, UV/vis and IR spectroscopies, mass spectrometry and elemental analysis. As for compound **IV-22**, the octahedral addition pattern was unambiguously confirmed by the high local symmetry of the C<sub>60</sub> core observed on the <sup>13</sup>C NMR spectrum of **IV-25**. The mixed [2:4] hexa-adducts **IV-25** with two defined addend zones (**Figure IV-16**) was obtained in 11 steps from 3-bromo-1-propanol in an overall yield of 5%.

Our methodology relies on the selective functionalization of two different reactive sites. By sequential CuAAC reactions, the post-functionalization of three different mixed C<sub>60</sub> hexa-adducts was performed in very good yields (75-99%).



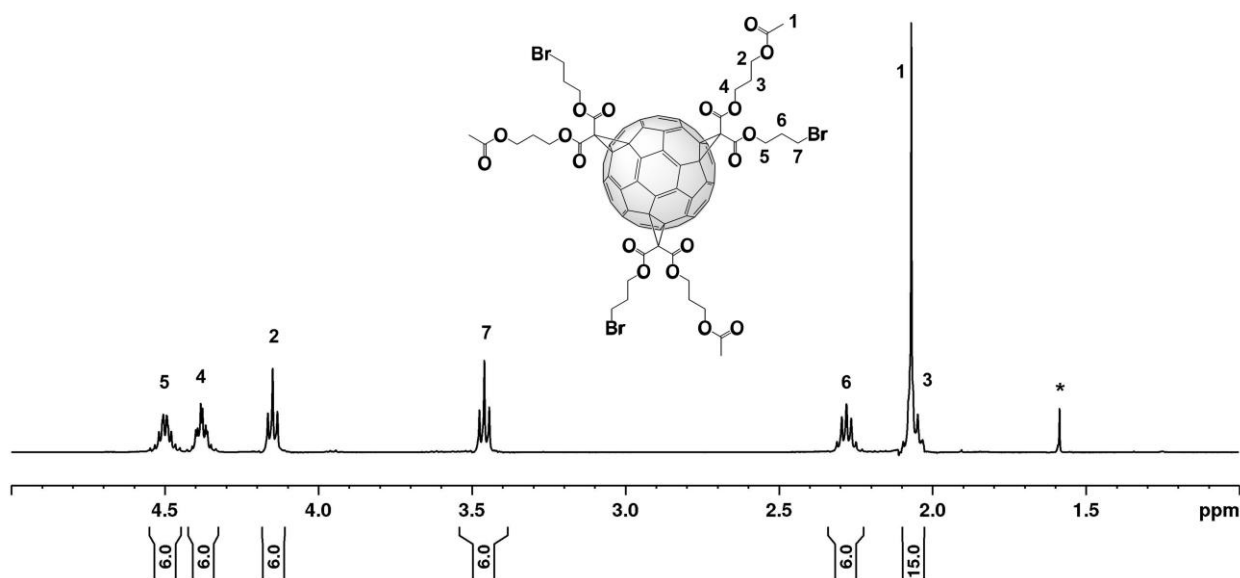
**Scheme IV-7.** Reagents and conditions: (i) Phenylacetylene, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (91%); (ii) NaN<sub>3</sub>, DMF, rt, 12 h (98%); (iii) 4-ethynylanisole, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (75%).



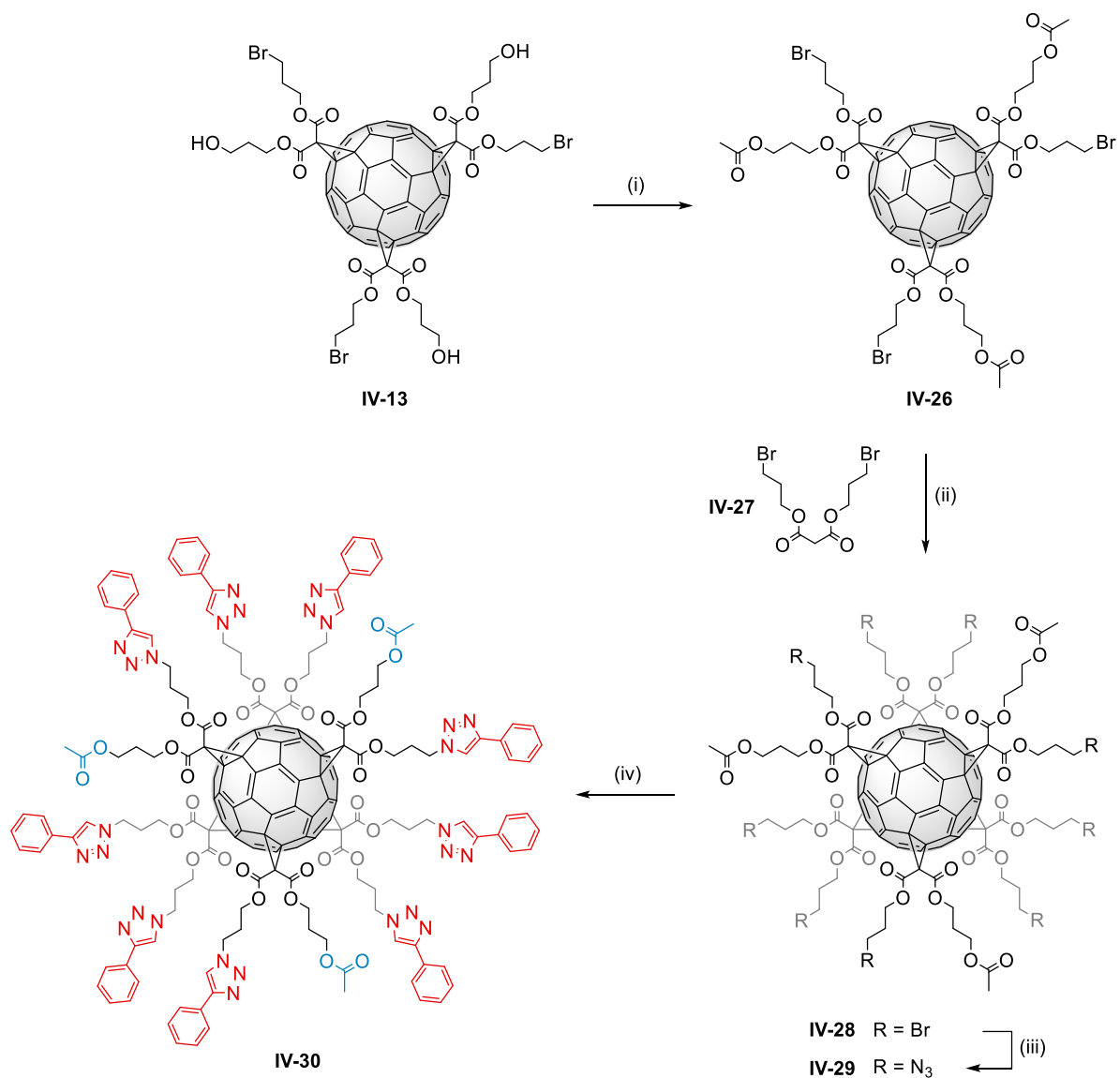
**Figure IV-16 :** Illustration of the two different addend zones of the mixed C<sub>60</sub> hexa-adduct IV-25.

### 3. Synthesis of mixed C<sub>60</sub> hexa-adducts by esterification and “click” reactions.

Based on the chemical reactivity of the alcohol function, tris-adduct triol **IV-13** can be post-functionalized first by an esterification reaction (Scheme IV-8). Treatment of **IV-13b** with acetyl chloride in the presence of pyridine gave tri-ester **IV-26** in 79% yield. As shown in [Figure IV-18](#), the <sup>1</sup>H NMR spectrum of **IV-26** reveals the disappearance of the CH<sub>2</sub>OH signals at δ = 3.6-3.7 ppm and the formation of the acetyl esters (at δ = 4.15 ppm for the CH<sub>2</sub>OOC signal and δ = 2.07 ppm for the OOCCH<sub>3</sub> signal). Reaction of **IV-26** with malonate **IV-27** (6 equiv.), CBr<sub>4</sub> (50 equiv.) and DBU (16 equiv.) in toluene at room temperature afforded hexa-adduct **IV-28**. Subsequent conversion of the bromide functions in azide by treatment with sodium azide followed by a CuAAC reaction with phenylacetylene gave the final mixed [3:3] hexa-adducts **IV-30**. Compound **IV-30** has been characterized by <sup>1</sup>H and <sup>13</sup>C NMR, UV/vis and IR spectroscopies, mass spectrometry and elemental analysis. As in the case of hexa-adducts **IV-22** and **IV-25**, the molecular symmetry (C<sub>3</sub>) of **IV-30** is deduced from the <sup>1</sup>H and <sup>13</sup>C NMR spectra and confirmed the octahedral addition pattern. As proof of the C<sub>3</sub>-symmetry, the <sup>13</sup>C NMR spectrum shows only one resonance for each C atoms of the side chains bearing the terminal acetyl group, as shown in [Figure IV-19](#). **IV-30** has a 3:9 function ratio regioselectively distributed ([Figure IV-17](#)).



[Figure IV-18](#) : <sup>1</sup>H NMR spectrum (400 MHz, CDCl<sub>3</sub>) of **IV-26**. (\*) water.



**Scheme IV-8.** Reagents and conditions: (i) Acetyl chloride, pyridine, THF, 0 °C (79%); (ii) **IV-32**, CBr<sub>4</sub>, DBU, PhMe, rt, 2 h (67%); (iii) NaN<sub>3</sub>, DMF, rt, 12 h (97%); (iv) Phenylacetylene, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (94%).

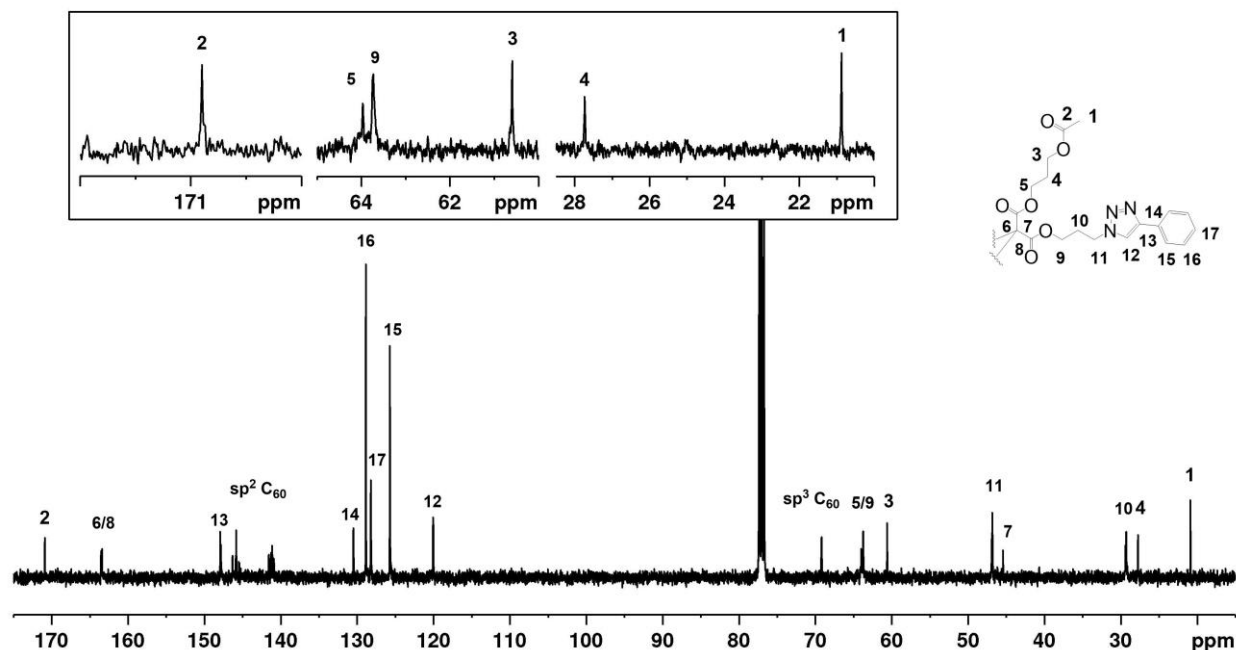


Figure IV-19:  $^{13}\text{C}$  NMR spectrum (100 MHz,  $\text{CDCl}_3$ ) of **IV-30** showing its threefold symmetry. Inset: detailed view showing one resonance for each C atoms of the side chains bearing the acetyl.

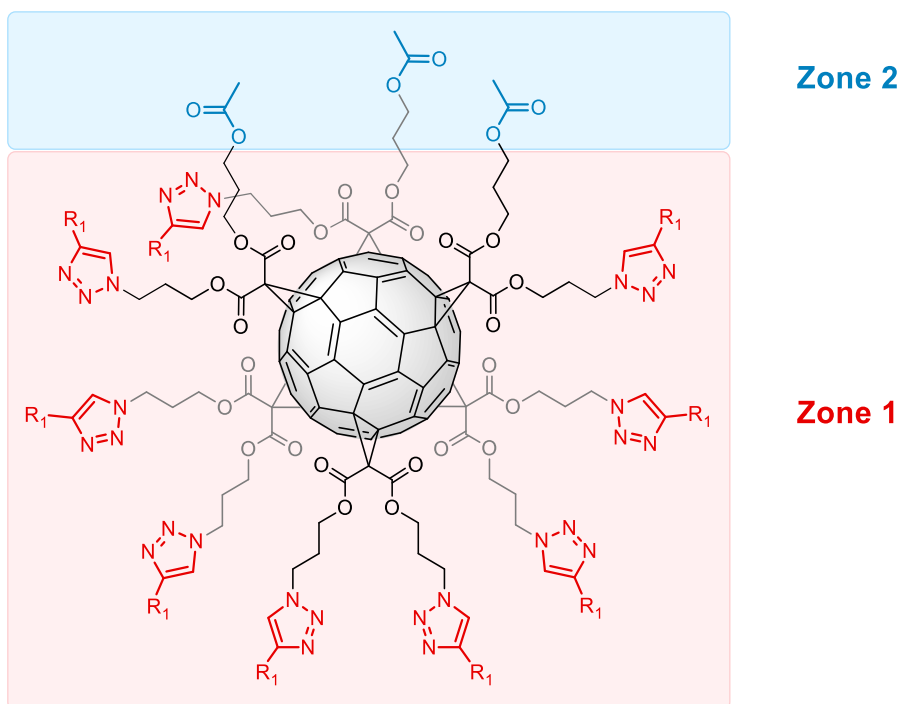
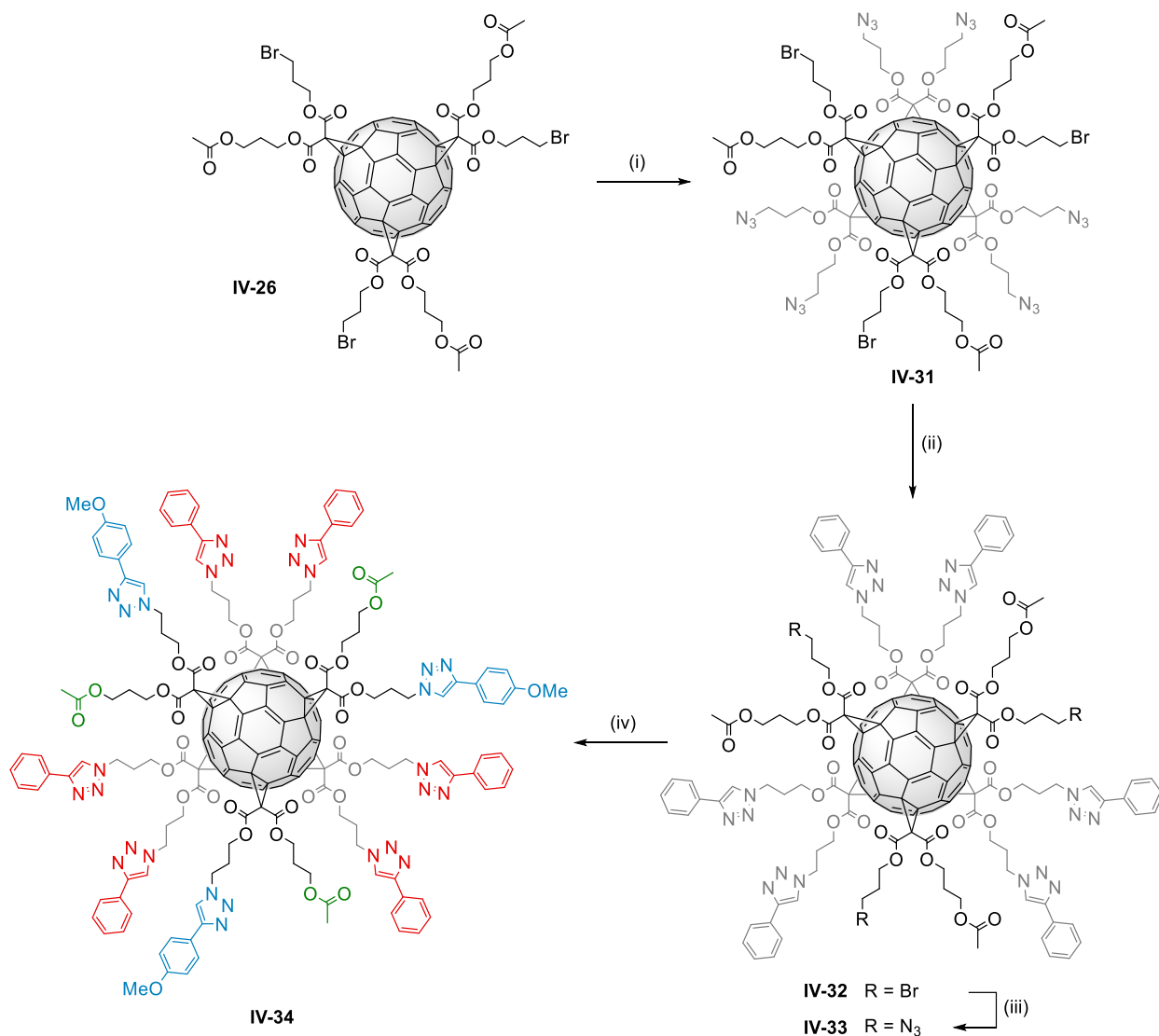


Figure IV-20 : Illustration of the two different addend zones of the mixed  $\text{C}_{60}$  hexa-adduct **IV-30**.

As shown in **Scheme IV-9**, the preparation of the mixed [3:3] hexa-adduct **IV-34** was performed from the tri-ester **IV-26**. The use of malonate **IV-12** instead of the malonate **IV-27** provided a third different reactive site. Treatment of **IV-26** with **IV-12** under modified Bingel conditions afforded hexa-adduct **IV-31** in 73% yield. Subsequent CuAAC reaction with phenylacetylene gave the first “clicked” derivative (**IV-32**) in an excellent yield (99%). Finally,

conversion of the bromide functions into azides followed by a second CuAAC reaction with 4-ethynylanisole gave the final mixed [3:3] C<sub>60</sub> hexa-adducts **IV-34** with a (3:3:6) function ratio. As for compound **IV-30**, the chemical structure of **IV-34** was confirmed by <sup>1</sup>H and <sup>13</sup>C NMR, UV/vis and IR spectroscopies, mass spectrometry and elemental analysis. In the case of **IV-34**, the three different addends zones have been selectively post-functionalized (Figure IV-21).



**Scheme IV-9.** Reagents and conditions: (i) **IV-12**, CBr<sub>4</sub>, DBU, PhMe, rt, 2 h (73%); (ii) Phenylacetylene, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (99%); (iii) NaN<sub>3</sub>, DMF, rt, 12 h (94%); (iv) 4-ethynylanisole, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (90%).



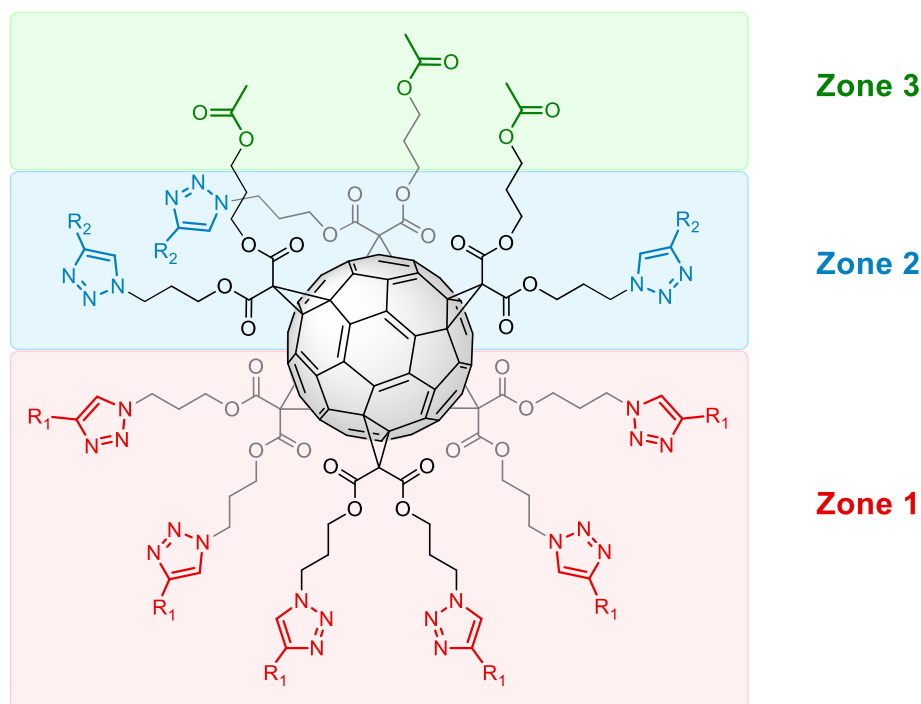


Figure IV-21 : Illustration of the three different addend zones of the mixed hexa-adduct **IV-34**.

## 4. Conclusion.

Based on our efficient methodologies for the preparation of *e,e,e* C<sub>60</sub> tris-adducts and *e* C<sub>60</sub> bis-adducts, an easy access to mixed C<sub>60</sub> hexa-adducts building blocks was possible. The modification of the side chains of the malonate addends has provided various functionalized mixed C<sub>60</sub> hexa-adducts. The sequential CuAAC reactions has been efficient for the post-functionalization of the C<sub>60</sub> hexa-adducts. Moreover, the utilization of an esterification reaction has diversified the scope of the post-functionalization possibilities and two more mixed hexa-adducts were obtained.

With the control of the regiochemistry and the stereochemistry of our methodology, up to three different addend zones can be defined according to the mixed C<sub>60</sub> hexa-adduct. These different addend zones have been selectively post-functionalized. The high tolerance of the CuAAC reactions to a wide variety of functions will thus provide a possible access to unprecedented globular multifunctional nanomaterials with a controlled distribution of functional groups on the spherical framework.

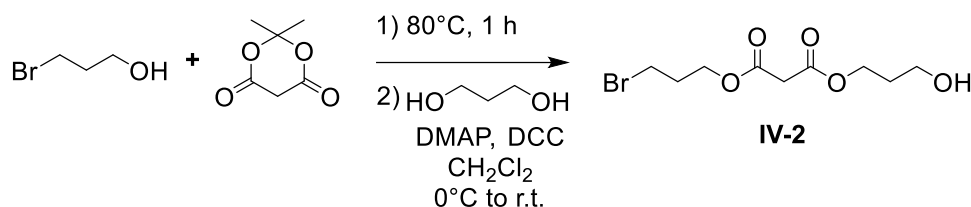
## 5. Bibliography.

- [1] A. Hirsch, O. Vostrowsky, *Eur. J. Org. Chem.* **2001**, 2001, 829–848.
- [2] H. Li, S. A. Haque, A. Kitaygorodskiy, M. J. Meziani, M. Torres-Castillo, Y.-P. Sun, *Org. Lett.* **2006**, 8, 5641–5643.
- [3] F. Djojo, E. Ravanelli, O. Vostrowsky, A. Hirsch, *Eur. J. Org. Chem.* **2000**, 2000, 1051–1059.
- [4] A. Herzog, A. Hirsch, O. Vostrowsky, *Eur. J. Org. Chem.* **2000**, 2000, 171–180.
- [5] B. Kräutler, T. Müller, J. Maynollo, K. Gruber, C. Kratky, P. Ochsenbein, D. Schwarzenbach, H.-B. Bürgi, *Angew. Chem. Int. Ed. Engl.* **1996**, 35, 1204–1206.
- [6] R. Schwenninger, T. Müller, B. Kräutler, *J. Am. Chem. Soc.* **1997**, 119, 9317–9318.
- [7] U. Reuther, T. Brandmüller, W. Donaubaue, F. Hampel, A. Hirsch, *Chem. – Eur. J.* **2002**, 8, 2261–2273.
- [8] M. Braun, U. Hartnagel, E. Ravanelli, B. Schade, C. Böttcher, O. Vostrowsky, A. Hirsch, *Eur. J. Org. Chem.* **2004**, 2004, 1983–2001.
- [9] C. P. Ioannou, G. I. Ioannou, E. E. Moushi, K. Velonia, N. Chronakis, *Eur. J. Org. Chem.* **2015**, 2015, 4598–4602.
- [10] F. Beuerle, A. Hirsch, *Chem. – Eur. J.* **2009**, 15, 7447–7455.
- [11] A. Gmehling, A. Hirsch, *Eur. J. Org. Chem.* **2013**, 2013, 5093–5105.
- [12] L. Isaacs, R. F. Haldimann, F. Diederich, *Angew. Chem. Int. Ed. Engl.* **1994**, 33, 2339–2342.
- [13] L. Isaacs, P. Seiler, F. Diederich, *Angew. Chem. Int. Ed. Engl.* **1995**, 34, 1466–1469.
- [14] R. F. Haldimann, F.-G. Klärner, F. Diederich, *Chem. Commun.* **1997**, 237–238.
- [15] L. Isaacs, F. Diederich, R. F. Haldimann, *Helv. Chim. Acta* **1997**, 80, 317–342.
- [16] N. Tirelli, F. Cardullo, T. Habicher, U. W. Suter, F. Diederich, *J. Chem. Soc. Perkin Trans. 2* **2000**, 193–198.
- [17] W. Qian, Y. Rubin, *Angew. Chem. Int. Ed.* **1999**, 38, 2356–2360.
- [18] W. Qian, Y. Rubin, *Angew. Chem. Int. Ed.* **2000**, 39, 3133–3137.
- [19] F. Beuerle, A. Hirsch, *Chem. – Eur. J.* **2009**, 15, 7434–7446.
- [20] I. Nierengarten, J.-F. Nierengarten, *Chem. Rec.* **2015**, 15, 31–51.
- [21] J. Iehl, R. P. de Freitas, J.-F. Nierengarten, *Tetrahedron Lett.* **2008**, 49, 4063–4066.
- [22] J. Iehl, R. P. de Freitas, B. Delavaux-Nicot, J.-F. Nierengarten, *Chem. Commun.* **2008**, 2450–2452.

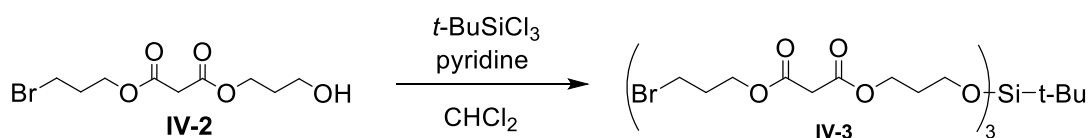
## 6. Experimental part.

**General:** Reagents and solvents were purchased as reagent grade and used without further purification. All reactions were performed in standard glassware under an inert Ar atmosphere. Evaporation and concentration were done at water aspirator pressure and drying in vacuo at  $10^{-2}$  Torr. Column chromatography: silica gel 60 (230-400 mesh, 0.040-0.063 mm) was purchased from E. Merck. Thin Layer Chromatography (TLC) was performed on aluminum sheets coated with silica gel 60 F<sub>254</sub> purchased from E. Merck. IR spectra ( $\text{cm}^{-1}$ ) were recorded on a Perkin–Elmer Spectrum One Spectrophotometer. UV/Visible spectra were recorded on a Perkin-Elmer Lambda 35 UV/Vis Spectrometer. NMR spectra were recorded on a Bruker AC 300 or AC 400 with solvent peaks as reference at 25°C. MALDI-TOF-MS were obtained on a Bruker ULTRAFLEX TOF/TOF mass spectrometer with a dithranol matrix.

### Synthesis.

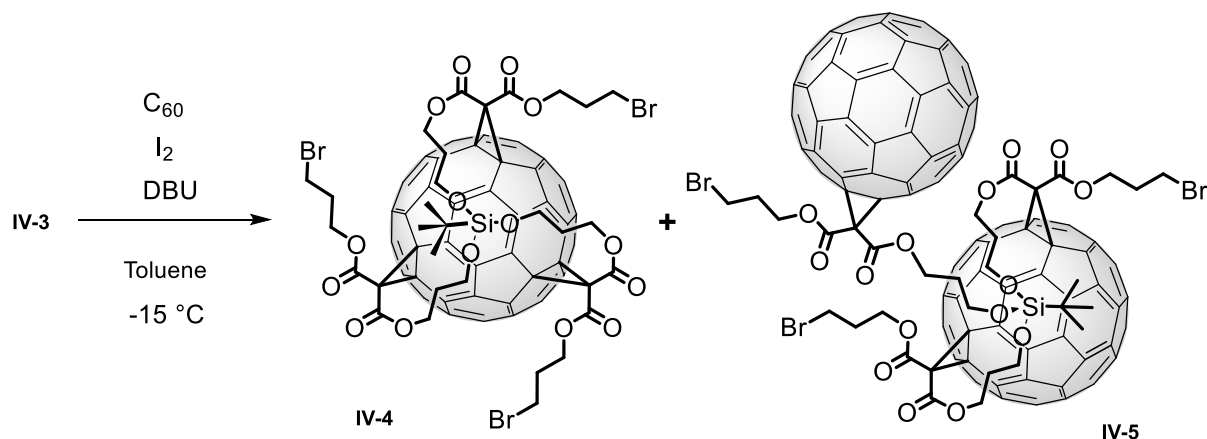


**Compound IV-2:** 3-Bromopropanol (4.16 mL, 46 mmol) and 2,2-dimethyl-1,3-dioxane-4,6-dione (6.63 g, 46 mmol) were stirred at 80°C. After 1 h the mixture was dried under reduced pressure and used in the next step without further purification. DCC (10.9 g, 53 mmol) was added to a solution of the crude, 3-propanediol (10.0 mL, 138 mmol) and DMAP (1.12 g, 9.2 mmol) in  $\text{CH}_2\text{Cl}_2$  (110 mL) at 0°C. The resulting mixture was stirred overnight at room temperature, then filtered on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 8:2) and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{EtOAc}$ , 9:1) yielded **IV-2** (6.08 g, 21.5 mmol, 46%) as a colorless oil. IR (neat):  $\nu = 3451$  (br, OH), 1731 ( $\text{C}=\text{O}$ )  $\text{cm}^{-1}$ .  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 4.33$  (t,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{OOC}$ ), 4.31 (t,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{OOC}$ ), 3.73 (t,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{OH}$ ), 3.47 (t,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ), 3.41 (s, 2 H,  $\text{OOCCH}_2\text{COO}$ ), 2.21 (quint.,  $^3J = 6$  Hz, 2 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{Br}$ ), 1.91 (quint.,  $^3J = 6$  Hz, 2 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{OH}$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 75 MHz):  $\delta = 166.7, 166.4, 63.3, 62.7, 59.2, 41.4, 31.5, 31.4, 29.1$  ppm.



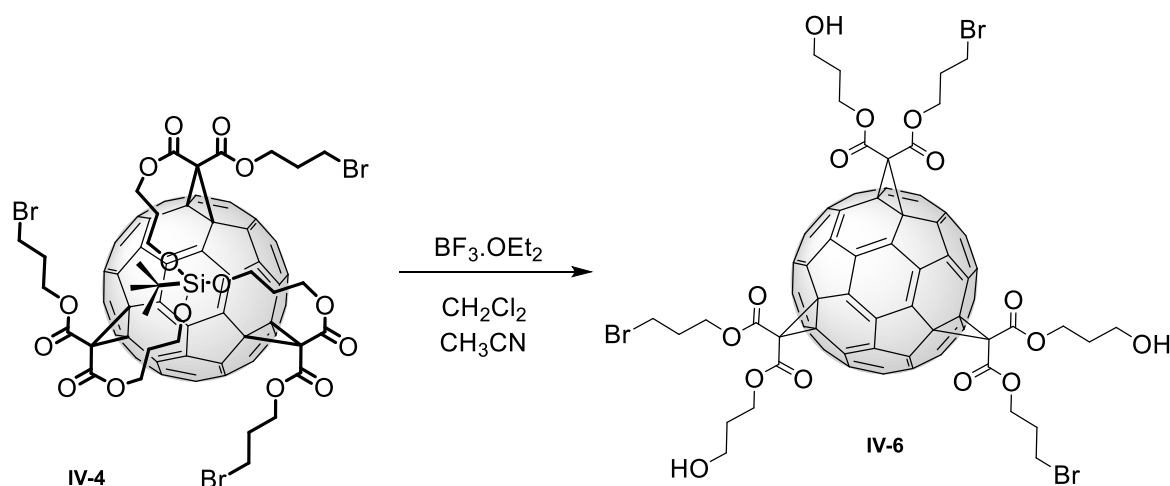
**Compound IV-3:**  $t\text{-BuSiCl}_3$  (47 mg, 0.25 mmol) was added to a solution of **IV-2** (210 mg, 0.74 mmol) and pyridine (0.07 mL, 0.8 mmol) in  $\text{CH}_2\text{Cl}_2$  (4 mL). After 18 h, the reaction mixture was filtered on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 97:3) and concentrated. Column chromatography on  $\text{SiO}_2$

(CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 99:1) yielded **IV-3** (151 mg, 0.16 mmol, 66%) as a colorless oil. IR (neat):  $\nu = 1732$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.29$  (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.27 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 3.85 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OSi), 3.46 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.39 (s, 6 H, OOCCH<sub>2</sub>COO), 2.19 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OSi), 1.90 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 0.94 (s, 9 H, *t*-BuSi) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.4$ , 166.3, 63.1, 62.5, 59.5, 41.4, 31.5, 29.1, 26.3, 17.7 ppm.

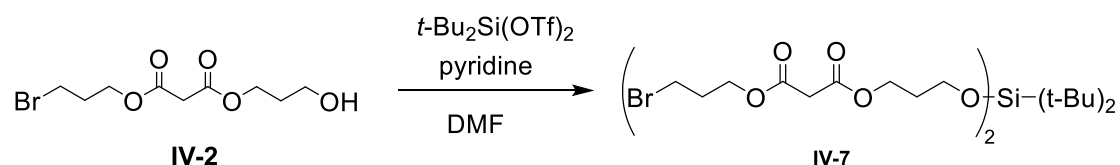


**Compound IV-4 & IV-5:** DBU (0.81 mL, 5.3 mmol) was added to a solution of C<sub>60</sub> (550 mg, 0.76 mmol), **IV-3** (710 mg, 0.76 mmol) and I<sub>2</sub> (639 mg, 2.5 mmol) in toluene (1.1 L) at -15 °C. The resulting mixture was stirred for 1 h at -15 °C, filtered on SiO<sub>2</sub> (cyclohexane to CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5 to 8:2) yielded **IV-4** (150 mg, 91 μmol, 12%) as a red glassy solid and **IV-5** (53 mg, 22 μmol, 3%) as a brown glassy solid. **IV-4:** IR (neat):  $\nu = 1740$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 233$  (1.3x10<sup>5</sup>), 251 (1.3x10<sup>5</sup>), 283 (9x10<sup>4</sup>), 302 (sh, 7x10<sup>4</sup>), 483 (8x10<sup>3</sup>), 563 (sh, 2x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta = 4.56$  (m, 3 H, CHH'CHH'OOC), 4.51 (m, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.24 (m, 3 H, CHH'CHH'OOC), 3.60 (m, 6 H, CHH'CH<sub>2</sub>OSi), 3.49 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.30 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 1.94 (m, 6 H, CHH'CHH'CH<sub>2</sub>OSi), 0.89 (s, 9 H, *t*-BuSi) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz):  $\delta = 163.3$ , 163.0, 147.0, 146.9, 146.8, 146.7, 146.4 (two peaks), 146.3, 146.1, 145.9, 144.4, 143.5, 143.4, 143.2, 143.1, 142.9, 142.5, 141.6, 141.0, 70.8, 70.1, 64.7, 64.0, 59.7, 52.7, 31.6, 31.2, 28.9, 26.1, 17.6 ppm. MS-MALDI-TOF : 1646.0 ([M]<sup>+</sup>, calcd for C<sub>91</sub>H<sub>45</sub>O<sub>15</sub>Br<sub>3</sub>Si: 1646.0). **IV-5:** IR (neat):  $\nu = 1745$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 256$  (1.5x10<sup>5</sup>), 325 (sh, 5x10<sup>4</sup>), 422 (3x10<sup>3</sup>), 425 (3x10<sup>3</sup>), 480 (3x10<sup>3</sup>), 553 (sh, 2x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.84$  (m, 1 H, CHH'CHH'OOC), 4.73 (m, 1 H, CHH'CHH'OOC), 4.68-4.55 (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.40 (m, 1 H, CHH'CHH'OOC), 4.30 (m, 1 H, CHH'CHH'OOC), 4.06 (m, 2 H, CH<sub>2</sub>CH<sub>2</sub>OSi), 3.88-3.74 (m, 4 H, CHH'CHH'OSi), 3.58 (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.55 (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.53 (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.40 (quint., <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 2.34 (m, 4 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 2.15 (quint., <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OSi), 1.99 (m, 4 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OSi), 0.91 (s, 9 H, *t*-BuSi) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.5$  (two peaks), 163.4, 163.2, 162.9, 148.3, 147.4, 147.3, 146.7, 146.6, 146.5, 146.4, 146.2, 146.1, 146.0, 145.7, 145.6, 145.3 (two peaks), 145.2 (three peaks), 145.1 (three peaks), 145.0 (two peaks), 144.9 (two peaks), 144.8,

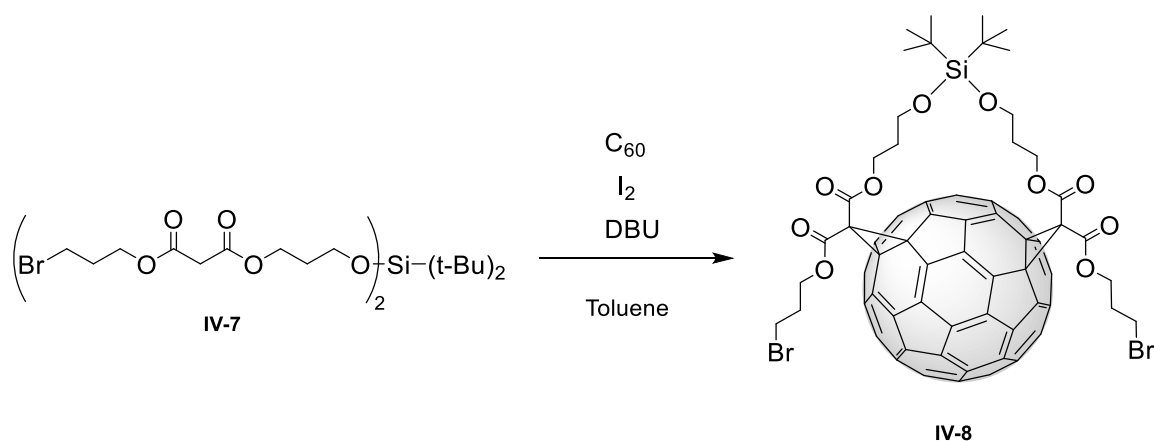
144.7 (four peaks), 144.6 (two peaks), 144.5, 144.3, 144.2, 144.1, 143.9 (two peaks), 143.8, 143.7 (two peaks), 143.6 (two peaks), 143.5 (two peaks), 143.1 (four peaks), 143.0 (two peaks), 142.3 (two peaks), 142.2, 142.1, 141.9 (three peaks), 141.7 (two peaks), 141.5, 141.0 (two peaks), 140.7, 139.6, 139.5, 139.3, 138.9, 138.8, 138.7, 71.7, 71.5 (three peaks), 70.4, 64.9 (two peaks), 64.8, 64.6, 63.8, 63.3, 60.0, 59.6, 58.6, 53.7, 52.1, 51.3, 32.1, 31.8, 31.5, 31.3, 29.1, 29.0 (two peaks), 26.3, 17.6 ppm. MS-MALDI-TOF : 2367.0 ([M+H]<sup>+</sup>, calcd for C<sub>151</sub>H<sub>46</sub>O<sub>15</sub>Br<sub>3</sub>Si : 2367.0).



**Compound IV-6:** BF<sub>3</sub>.OEt<sub>2</sub> (40 μL, 267 μmol) was added to a solution of IV-4 (44 mg, 27 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/MeCN (9/4 mL). The solution was stirred overnight at room temperature, then filtered on SiO<sub>2</sub> (THF) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 97:3) yielded IV-6 (36 mg, 23 μmol, 86%) as a red glassy solid. IR (neat):  $\nu$  = 3208 (br, OH), 1742 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon)$  = 235 (8x10<sup>4</sup>), 251 (9x10<sup>4</sup>), 283 (6x10<sup>4</sup>), 305 (sh, 4x10<sup>4</sup>), 396 (sh, 2x10<sup>3</sup>), 483 (4x10<sup>3</sup>), 523 (sh, 3x10<sup>3</sup>), 562 (sh, 1x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  = 4.58-4.52 (m, 3 H, CH<sub>2</sub>CHH'OOOC), 4.49-4.42 (m, 6 H, CHH'CHH'OOOC), 4.25-4.21 (m, 3 H, CHH'CHH'OOOC), 3.74-3.70 (m, 3 H, CHH'CHH'OH), 3.62-3.56 (m, 3 H, CHH'CHH'OH), 3.49 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.30 (quint., <sup>3</sup>J = 6 Hz, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 1.88 (m, 6 H, CHH'CHH'CHH') ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  = 163.1, 163.0, 147.2, 147.0, 146.9, 146.8, 146.4, 146.3, 146.1, 145.9, 145.8, 144.3, 143.5, 143.3, 143.2, 143.1, 142.7, 141.3, 140.9, 70.9, 70.0, 64.6, 62.6, 57.8, 52.6, 31.2, 28.9, 28.7 ppm. MS-MALDI-TOF : 1564.0 ([M]<sup>+</sup>, calcd for C<sub>87</sub>H<sub>39</sub>O<sub>15</sub>Br<sub>3</sub>: 1564.0).

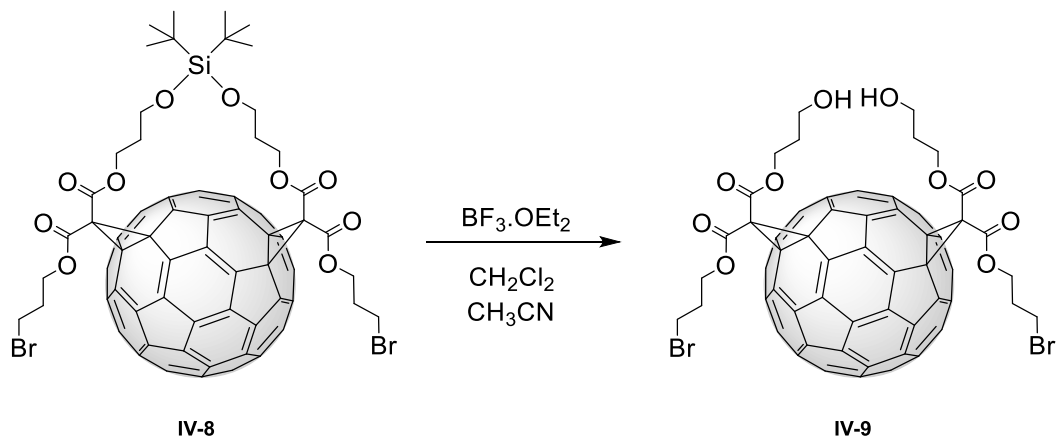


**Compound IV-7:** A solution of **IV-2** (3.53 g, 12.46 mmol), pyridine (1.26 mL, 15.5 mmol) and  $(t\text{-Bu})_2\text{Si(OTf)}_2$  (2.02 mL, 6.22 mmol) in anhydrous DMF (20 mL) was stirred at room temperature overnight. Then,  $\text{H}_2\text{O}$  was added and the aqueous layer extracted with  $\text{Et}_2\text{O}$ . The organic layer was washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 8:2 to  $\text{CH}_2\text{Cl}_2$ ) yielded **IV-7** (3.00 g, 4.22 mmol, 68%) as a colorless oil. IR (neat):  $\nu = 1735$  ( $\text{C}=\text{O}$ )  $\text{cm}^{-1}$ .  $^1\text{H NMR}$  ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 4.30$  (t,  $^3J = 6$  Hz, 8 H,  $\text{CH}_2\text{CH}_2\text{OOC}$ ), 3.91 (t,  $^3J = 6$  Hz, 4 H,  $\text{CH}_2\text{CH}_2\text{OSi}$ ), 3.46 (t,  $^3J = 6$  Hz, 4 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ), 3.39 (s, 4 H,  $\text{OOCCH}_2\text{COO}$ ), 2.20 (quint.,  $^3J = 6$  Hz, 4 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{Br}$ ), 1.91 (quint.,  $^3J = 6$  Hz, 4 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{OSi}$ ), 1.00 (s, 18 H,  $\text{Si(t-Bu)}_2$ ) ppm.  $^{13}\text{C NMR}$  ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 166.4, 166.3, 63.1, 62.7, 60.1, 41.4, 31.7, 31.5, 29.1, 27.8, 21.2$  ppm.

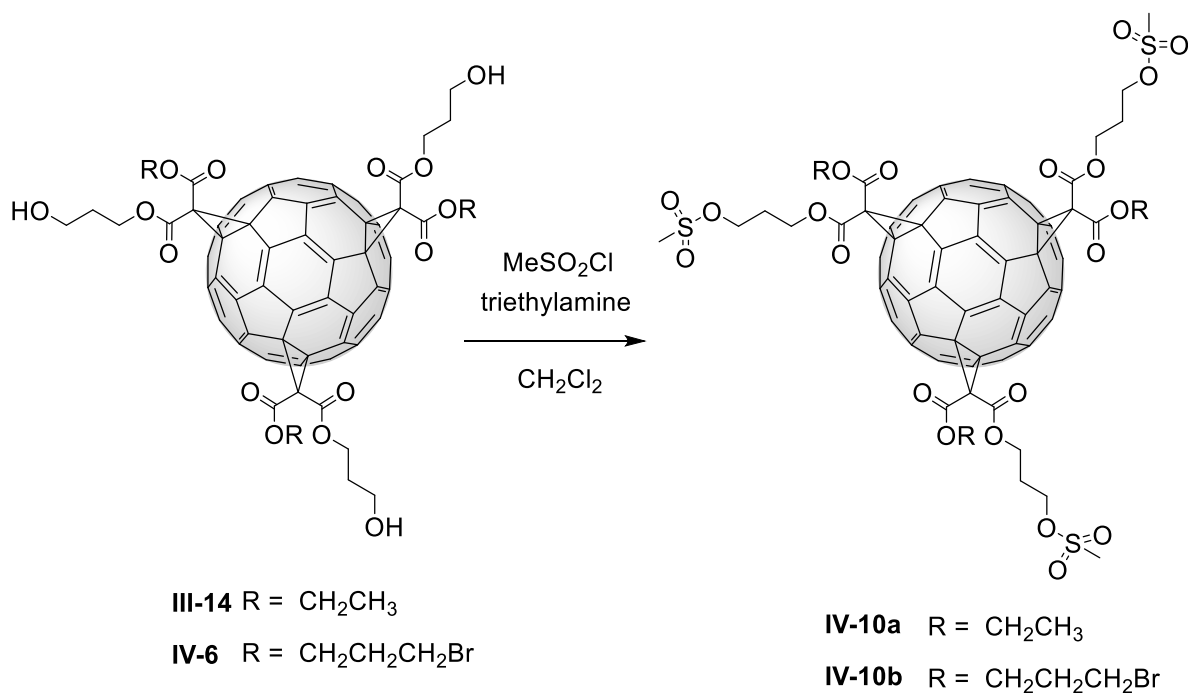


**Compound IV-8:** DBU (0.73 mL, 4.9 mmol) was added to a solution of  $\text{C}_{60}$  (700 mg, 0.97 mmol), **IV-7** (690 mg, 0.97 mmol) and  $\text{I}_2$  (616 mg, 2.4 mmol) in toluene (1.4 L) at room temperature. The resulting mixture was stirred for 1 h, directly filtered on  $\text{SiO}_2$  ( $\text{cyclohexane}$  then  $\text{CH}_2\text{Cl}_2$ ) and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{cyclohexane}$ , 5:5) yielded **IV-8** (660 mg, 0.46 mmol, 48%) as a brown glassy solid. IR (neat):  $\nu = 1747$  ( $\text{C}=\text{O}$ )  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 252$  ( $1.4 \times 10^5$ ), 313 (sh,  $5 \times 10^4$ ), 397 ( $5 \times 10^3$ ), 409 (sh,  $3 \times 10^3$ ), 421 ( $3 \times 10^3$ ), 477 ( $4 \times 10^3$ ) nm.  $^1\text{H NMR}$  ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 4.75$  (m, 2 H), 4.60 (m, 4 H), 4.41 (m, 2 H), 3.89 (m, 2 H), 3.81 (m, 2 H), 3.54 (t,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ), 3.53 (t,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ), 2.35 (m, 4 H), 1.99 (m, 4H), 1.02 (s, 9H,  $t\text{-BuSi}$ ), 0.90 (s, 9H,  $t\text{-BuSi}$ ) ppm.  $^{13}\text{C NMR}$  ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 164.1, 163.4, 163.2, 163.1, 148.5, 147.4, 147.3, 146.8, 146.6, 146.5$  (two peaks), 146.3, 146.2 (two peaks), 146.1, 145.8, 145.6, 145.3, 145.2, 145.1, 145.0, 144.9, 144.8, 144.7 (three peaks), 144.6, 144.5, 144.4, 144.2, 144.1, 144.0, 143.9, 143.8, 143.7 (two peaks), 143.6, 143.5 (two peaks), 143.2, 143.1 (two peaks), 142.3, 142.1, 142.0 (three peaks), 141.7, 141.5, 141.4, 140.6, 139.4 (two peaks), 138.8, 138.7, 71.6, 71.5 (two peaks), 70.3, 64.8, 64.4, 63.5, 60.3, 59.7, 53.9,

51.3, 32.2, 31.7, 31.4, 31.3, 29.0, 28.8, 27.8, 21.3 ppm. MS-MALDI-TOF: 1423.0 ( $[M+H]^+$ , calcd for  $C_{86}H_{43}O_{10}Br_2Si$ : 1423.1).



**Compound IV-9:** A solution of **IV-8** (1.32 g, 0.93 mmol) and  $BF_3 \cdot OEt_2$  (1.53 mL, 12.0 mmol) in  $CH_2Cl_2/CH_3CN$  (40/25 mL) was stirred overnight at room temperature, then filtered on  $SiO_2$  ( $CH_2Cl_2/MeOH$ , 95 :5) and concentrated. Column chromatography on  $SiO_2$  ( $CH_2Cl_2$  to  $CH_2Cl_2/MeOH$ , 97:3) yielded **IV-9** (1.08 g, 0.85 mmol, 91%) as a brown glassy solid. IR (neat):  $\nu = 3405$  (br, OH),  $1742$  (C=O)  $cm^{-1}$ . UV/Vis ( $CH_2Cl_2$ ):  $\lambda_{max}(\epsilon) = 252$  ( $1 \times 10^5$ ),  $313$  (sh,  $5 \times 10^4$ ),  $397$  ( $6 \times 10^3$ ),  $409$  (sh,  $4 \times 10^3$ ),  $421$  ( $4 \times 10^3$ ),  $481$  ( $5 \times 10^3$ ) nm.  $^1H$  NMR ( $CDCl_3$ , 400 MHz):  $\delta = 4.58$  (m, 8 H,  $CH_2CH_2OOC$ ),  $3.77$  (t,  $^3J = 6$  Hz, 2 H,  $CH_2CH_2OH$ ),  $3.76$  (t,  $^3J = 6$  Hz, 2 H,  $CH_2CH_2OH$ ),  $3.52$  (t,  $^3J = 6$  Hz, 4 H,  $CH_2CH_2Br$ ),  $2.34$  (m, 4 H,  $CH_2CH_2CH_2OH$ ),  $2.03$  (m, 4 H,  $CH_2CH_2CH_2Br$ ),  $1.95$  (br s, 2 H, OH) ppm.  $^{13}C$  NMR ( $CDCl_3$ , 100 MHz):  $\delta = 163.7, 163.5, 163.4, 163.3, 147.4$  (two peaks),  $147.3$  (two peaks),  $146.6, 146.5$  (two peaks),  $146.1, 146.0, 145.7, 145.6, 145.4, 145.1, 145.0, 144.8$  (two peaks),  $144.7$  (three peaks),  $144.4, 144.3, 144.1, 143.9$  (two peaks),  $143.8, 143.7, 143.5$  (two peaks),  $143.2$  (two peaks),  $142.9, 142.8, 142.5, 142.3, 142.0$  (two peaks),  $141.7, 141.6, 141.5, 138.8, 138.7, 138.6, 71.5, 71.4$  (two peaks),  $70.2, 64.9, 64.8, 64.4$  (two peaks),  $59.0, 53.4, 51.2, 31.5, 31.4$  (three peaks),  $29.1$  ppm. MS-MALDI-TOF: 1281.9 ( $[M]^+$ , calcd for  $C_{78}H_{26}O_{10}Br_2$ : 1281.9).

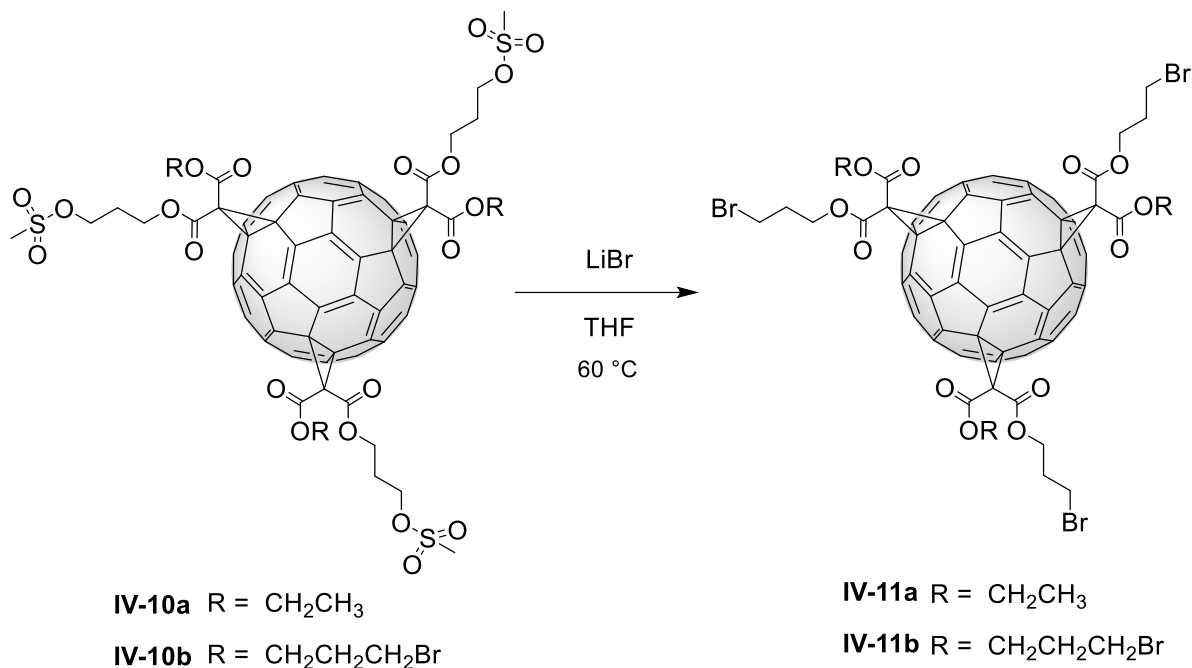


**Compound IV-10a:** MeSO<sub>2</sub>Cl (0.11 mL, 1.44 mmol) was added to a solution of **III-14** (311 mg, 0.24 mmol) and triethylamine (0.2 mL, 1.44 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (40 mL) at 0°C. After 2 h, the mixture was filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 99:1) yielded **IV-10a** (316 mg, 0.21 mmol, 86%) as a red glassy solid. IR (neat):  $\nu = 1740$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 237$  (1.2x10<sup>5</sup>), 251 (1.3x10<sup>5</sup>), 282 (9x10<sup>4</sup>), 305 (sh, 6x10<sup>4</sup>), 484 (7x10<sup>3</sup>), 524 (sh, 4x10<sup>3</sup>), 565 (sh, 2x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.43$  (m, 12 H, CH<sub>2</sub>OOC), 4.30 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OSO<sub>2</sub>CH<sub>3</sub>), 3.03 (s, 9 H, OSO<sub>2</sub>CH<sub>3</sub>), 2.17 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.37 (t, <sup>3</sup>J = 7Hz, 9 H, CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.2, 147.0, 146.8, 146.7, 146.6, 146.5, 146.4, 146.3, 145.7, 145.6, 144.9, 144.6, 144.4, 143.4, 142.7, 142.6, 141.8, 141.6, 141.0, 70.7, 69.9, 65.8, 63.4, 62.6, 52.6, 37.4, 28.4, 14.2$  ppm. MS-MALDI-TOF: 1518.2 ([M]<sup>+</sup>, calcd for C<sub>87</sub>H<sub>42</sub>O<sub>21</sub>S<sub>3</sub>: 1518.1).

**Compound IV-10b:** MeSO<sub>2</sub>Cl (16  $\mu$ L, 207  $\mu$ mol) was added to a solution of **IV-6** (36 mg, 23  $\mu$ mol) and triethylamine (29  $\mu$ L, 207  $\mu$ mol) in CH<sub>2</sub>Cl<sub>2</sub> (3 mL) at 0°C. After 30 min, the mixture was filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 97:3) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 99:1) yielded **IV-10b** (39 mg, 22  $\mu$ mol, 94%) as a red glassy solid. IR (neat):  $\nu = 1740$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 234$  (1x10<sup>5</sup>), 251 (1x10<sup>5</sup>), 283 (7x10<sup>4</sup>), 304 (sh, 5x10<sup>4</sup>), 483 (7x10<sup>3</sup>), 523 (sh, 4x10<sup>3</sup>), 564 (sh, 2x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta = 4.52$  (dt, <sup>2</sup>J = 1.5 Hz & <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.43 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.30 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OSO<sub>2</sub>CH<sub>3</sub>), 3.47 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.03 (s, 9 H, OSO<sub>2</sub>CH<sub>3</sub>), 2.29 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 2.18 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OSO<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 75 MHz):  $\delta = 163.1$  (two peaks), 147.0, 146.9, 146.7, 146.6, 146.5, 146.4, 145.7, 145.4, 145.3, 144.8, 144.5, 143.5, 142.7, 142.3, 141.5, 141.4, 141.1, 70.6, 69.8, 65.7,



64.8, 62.8, 52.6, 37.5, 31.2, 29.1, 28.4 ppm. MS-MALDI-TOF: 1797.9 (100%, [M]<sup>+</sup>, calcd for C<sub>90</sub>H<sub>45</sub>O<sub>21</sub>S<sub>3</sub>Br<sub>3</sub>: 1797.9), 1702.9 (12%, [M-CH<sub>2</sub>Br]<sup>+</sup>, calcd for C<sub>89</sub>H<sub>43</sub>O<sub>21</sub>S<sub>3</sub>Br<sub>2</sub>: 1703.0).

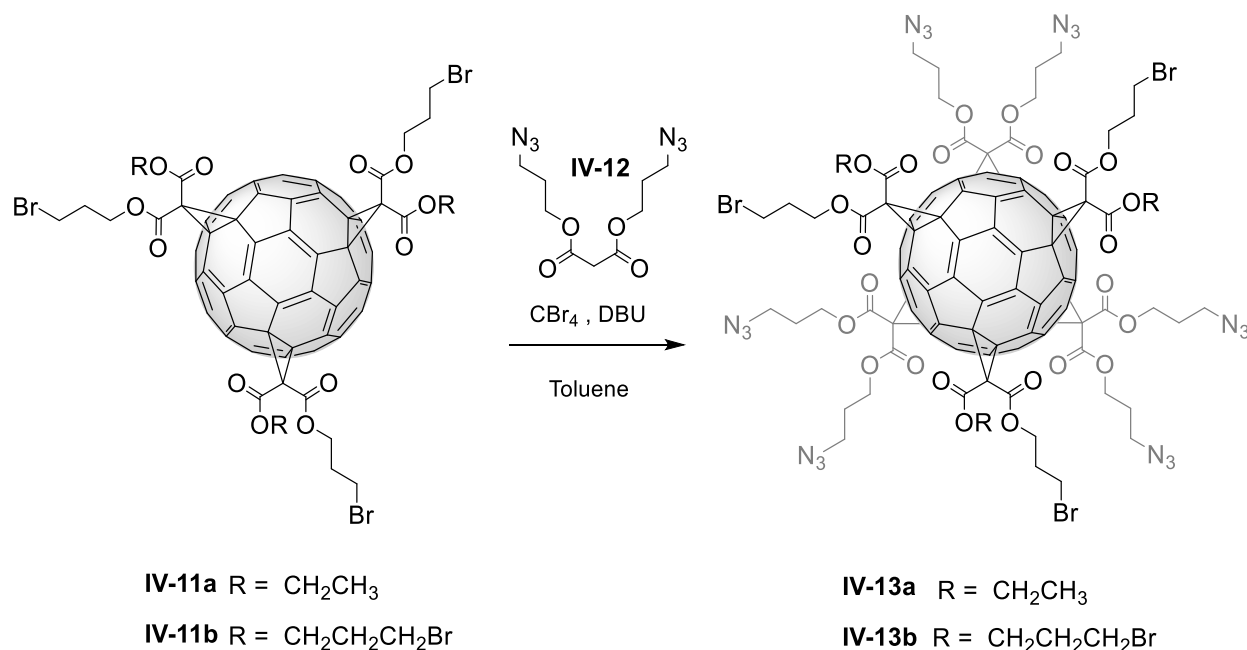


**Compound IV-11a:** A solution of **IV-10a** (304 mg, 0.20 mmol) and LiBr (104 mg, 1.2 mmol) was stirred overnight at 60°C, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 6:4) yielded **IV-11a** (244 mg, 0.16 mmol, 83%) as a red glassy solid. IR (neat):  $\nu = 1743$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 235$  (1.1x10<sup>5</sup>), 251 (1.1x10<sup>5</sup>), 282 (8x10<sup>4</sup>), 303 (sh, 6x10<sup>4</sup>), 484 (7x10<sup>3</sup>), 525 (sh, 4x10<sup>3</sup>), 564 (sh, 2x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.42$  (m, 12 H, CH<sub>2</sub>OOC), 3.46 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.25 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.38 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.3, 163.2, 147.1, 146.8, 146.7, 146.6, 146.5$  (two peaks), 146.3, 145.7, 145.5, 144.9, 144.8, 144.3, 143.4, 142.7, 142.5, 141.9, 141.7, 141.0, 70.8, 70.0, 64.5, 63.3, 52.6, 31.3, 28.9, 14.2 ppm. MS-MALDI-TOF: 1474.0 ([M]<sup>+</sup>, calcd for C<sub>84</sub>H<sub>33</sub>O<sub>12</sub>Br<sub>3</sub>: 1474.0).

**Compound IV-11b:** A solution of **IV-10b** (192 mg, 107 μmol) and LiBr (56 mg, 640 μmol) in THF (10 mL) was stirred overnight at 60°C, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) yielded **IV-11b** (184 mg, 105 μmol, 98%) as a red glassy solid. IR (neat):  $\nu = 1741$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 235$  (1.1x10<sup>5</sup>), 251 (1.1x10<sup>5</sup>), 282 (8x10<sup>4</sup>), 305 (sh, 6x10<sup>4</sup>), 484 (6x10<sup>3</sup>), 524 (sh, 4x10<sup>3</sup>), 565 (sh, 2x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.50$  (m, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.45 (m, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 3.47 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.45 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.29 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 2.26 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.1$  (two peaks), 147.1, 146.9, 146.7, 146.6 (two peaks), 146.5, 146.3, 145.7, 145.3, 145.2, 144.9, 144.5, 143.5, 142.7, 142.3, 141.6, 141.3, 141.0, 70.7, 69.8, 64.7 (two peaks), 52.6, 31.3, 31.2, 29.0 ppm. MS-

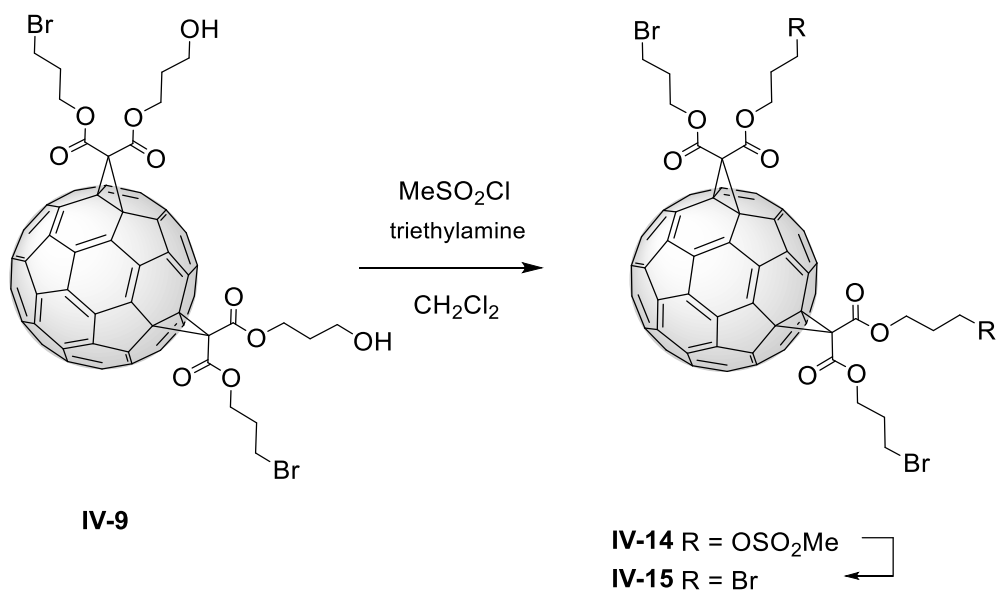
MALDI-TOF: 1751.8 (100%, [M]<sup>+</sup>, calcd for C<sub>87</sub>H<sub>36</sub>O<sub>12</sub>Br<sub>6</sub>: 1751.7), 1614.8 (16%, [M-(OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br)]<sup>+</sup>, calcd for C<sub>84</sub>H<sub>30</sub>O<sub>11</sub>Br<sub>5</sub>: 1614.8).

**Compound IV-12:** IV-12 was prepared as described in *Chem. Commun.* **2008**, 2450–2452.



**Compound IV-13a:** DBU (0.43 mL, 2.9 mmol) was added to a solution of **IV-2** (240 mg, 0.16 mmol), **IV-3** (220 mg, 0.81 mmol) and CBr<sub>4</sub> (2.12 g, 6.4 mmol) in toluene (35 mL). The mixture was stirred overnight at room temperature, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 98:2) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 96:4) followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) yielded **IV-4** (155 mg, 68 μmol, 42%) as an orange glassy solid. IR (neat):  $\nu = 2099$  (N<sub>3</sub>), 1742 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.42$  (t, <sup>3</sup>J = 6 Hz, 6 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 4.36 (t, <sup>3</sup>J = 6 Hz, 12 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 4.34 (q, <sup>3</sup>J = 7 Hz, 6 H, COOCH<sub>2</sub>CH<sub>3</sub>), 3.44 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.39 (t, <sup>3</sup>J = 6 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 2.24 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 1.96 (quint., <sup>3</sup>J = 6 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 1.34 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>2</sub>CH<sub>3</sub>) ppm.

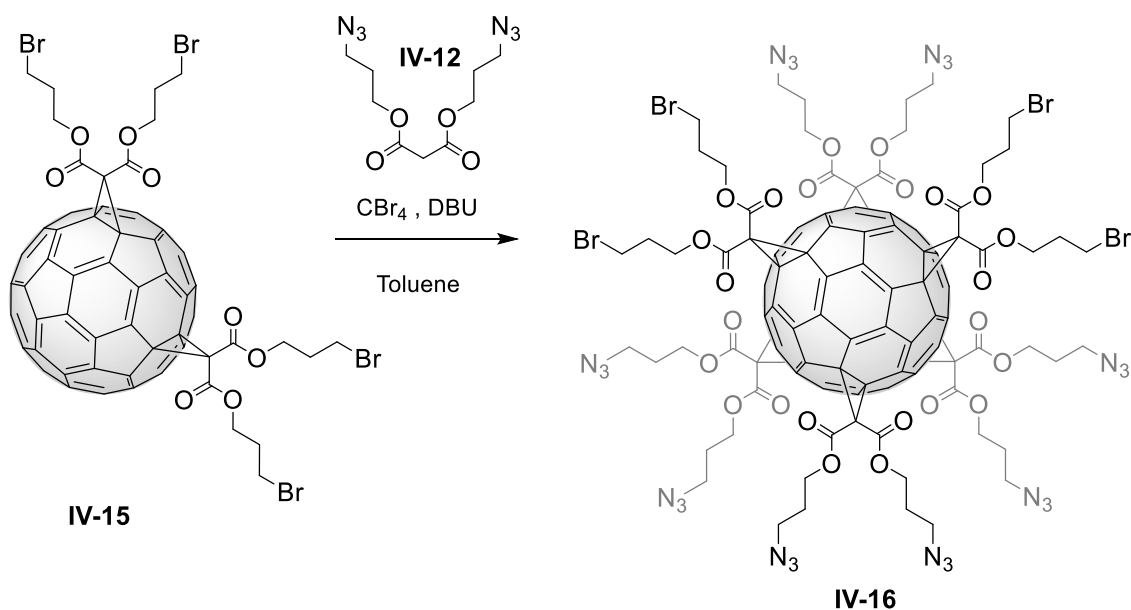
**Compound IV-13b:** DBU (0.14 mL, 0.9 mmol) was added to a solution of **IV-11b** (102 mg, 58 μmol), **IV-12** (94 mg, 0.35 mmol) and CBr<sub>4</sub> (965 mg, 2.9 mmol) in toluene (20 mL). The resulting mixture was stirred overnight at room temperature, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 98:2) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 98:2) yielded **IV-13b** (94 mg, 37 μmol, 63%) as an orange glassy solid. IR (neat):  $\nu = 2097$  (N<sub>3</sub>), 1741 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta = 4.43$  (t, <sup>3</sup>J = 6 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.37 (t, <sup>3</sup>J = 6 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 3.44 (t, <sup>3</sup>J = 6 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 3.40 (t, <sup>3</sup>J = 6 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.25 (quint., <sup>3</sup>J = 6 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 1.97 (quint., <sup>3</sup>J = 6 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>) ppm.



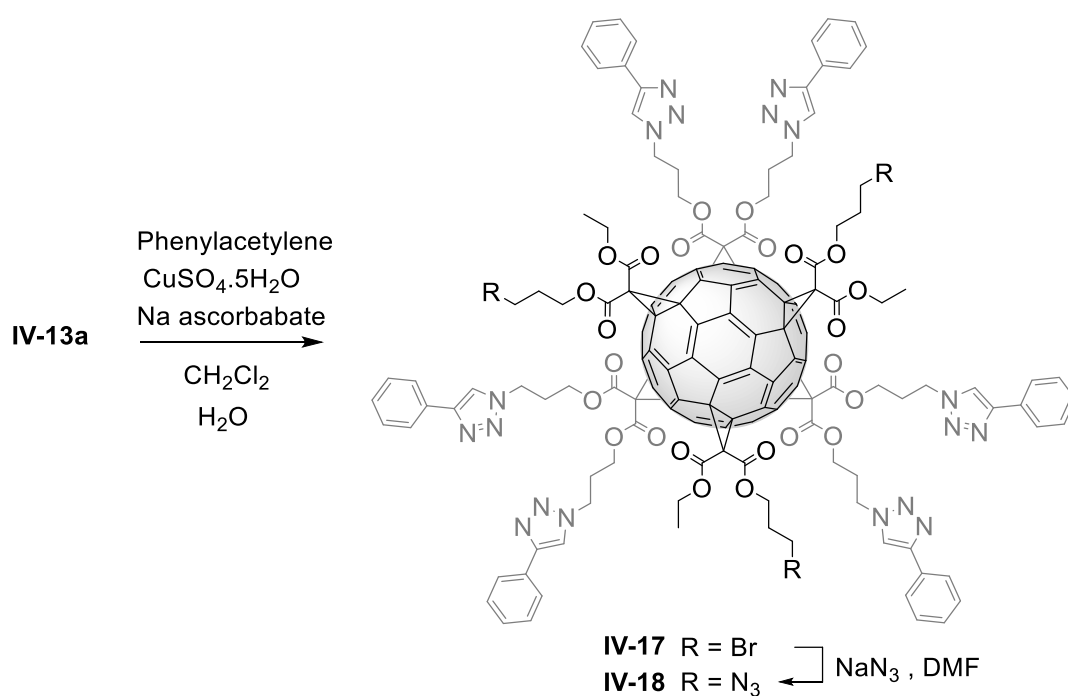
**Compound IV-14:** Methane sulfonyl chloride (0.27 mL, 3.5 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (2 mL) was added slowly to a solution of **IV-9** (1.30 g, 1.0 mmol) and triethylamine (0.56 mL, 4.0 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (145 mL) at 0°C. The mixture was stirred at 0°C for 15 min, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 9:1) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 97:3) yielded **IV-14** (1.37 g, 0.95 mmol, 93%) as a brown glassy solid. IR (neat):  $\nu = 1739$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 252$  (1.4x10<sup>5</sup>), 312 (sh, 5x10<sup>4</sup>), 397 (5x10<sup>3</sup>), 408 (sh, 3x10<sup>3</sup>), 421 (3x10<sup>3</sup>), 480 (4x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.58$  (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.37 (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>OS), 4.35 (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>OS), 3.51 (t, <sup>3</sup>J = 6 Hz, 4 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.06 (s, 3 H, OOSCH<sub>3</sub>), 3.05 (s, 3 H, OOSCH<sub>3</sub>), 2.34 (m, 4 H), 2.25 (m, 4 H) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.3$  (two peaks), 163.2 (two peaks), 147.4, 147.2, 146.6 (two peaks), 146.5, 146.1 (two peaks), 146.0, 145.7, 145.5, 145.4, 145.1, 145.0, 144.9, 144.8 (two peaks), 144.7, 144.4, 144.1 (two peaks), 143.8 (two peaks), 143.7 (two peaks), 143.6 (two peaks), 143.2, 143.1, 143.0 (two peaks), 142.8, 142.5, 142.4, 142.1, 141.7, 141.5 (two peaks), 138.8, 138.7 (two peaks), 71.4, 71.3, 70.1, 65.7 (two peaks), 65.0, 64.9, 63.0, 53.2, 51.0, 37.5, 31.3, 29.1, 28.5 (two peaks) ppm. MS-MALDI-TOF: 1437.9 ([M]<sup>+</sup>, calcd for C<sub>80</sub>H<sub>30</sub>O<sub>14</sub>S<sub>2</sub>Br<sub>2</sub>: 1437.9).

**Compound IV-15:** A solution of **IV-25** (1.29 g, 0.89 mmol) and LiBr (0.31 g, 3.6 mmol) in THF (40 mL) was stirred at 60°C for 21 h. Then the mixture was filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 6:4) yielded **IV-15** (1.21 g, 0.86 mmol, 96%) as a brown glassy solid. IR (neat):  $\nu = 1745$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 252$  (2x10<sup>5</sup>), 313 (sh, 5x10<sup>4</sup>), 397 (9x10<sup>3</sup>), 409 (sh, 6x10<sup>3</sup>), 421 (5x10<sup>3</sup>), 482 (6x10<sup>3</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.57$  (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 3.52 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.50 (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.34 (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.3$  (two peaks), 163.2, 147.4, 146.6, 146.5, 146.1 (two peaks), 145.7, 145.4, 145.1, 145.0, 144.9, 144.8, 144.7, 144.5, 144.2, 144.1, 143.8 (two peaks), 143.6, 143.2, 143.0, 142.5,

142.1, 141.6 (two peaks), 138.8, 138.7, 71.4 (two peaks), 70.1, 64.9 (two peaks), 53.2, 51.0, 31.4, 31.3, 29.0 ppm. MS-MALDI-TOF: 1408.2 ( $[M]^+$ , calcd for  $C_{78}H_{24}O_8Br_4$ : 1407.8).



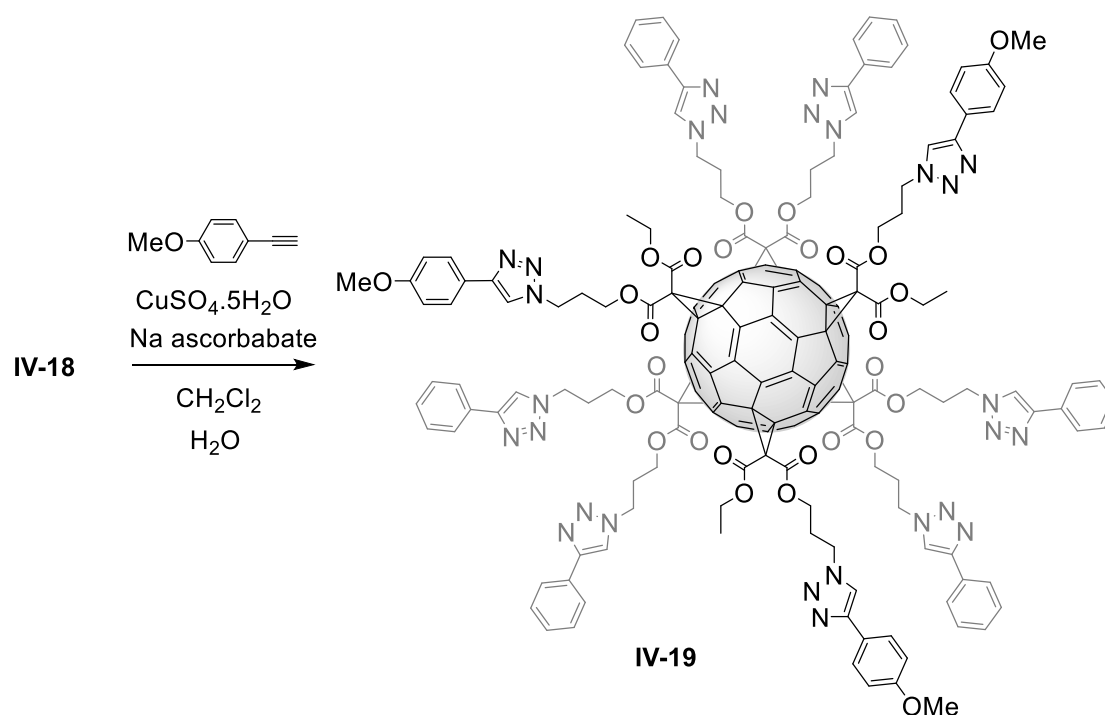
**Compound IV-16:** DBU (0.57 mL, 3.8 mmol) was added to a solution of IV-15 (270 mg, 0.19 mmol), IV-12 (311 mg, 1.15 mmol) and CBr<sub>4</sub> (3.18 g, 9.6 mmol) in toluene (45 mL). After 22 h, the mixture was filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 95:5) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 8:2 to CH<sub>2</sub>Cl<sub>2</sub>/EtOAc, 96:4) yielded IV-16 (267 mg, 0.11 mmol, 56%) as an orange glassy solid. IR (neat):  $\nu = 2097$  (N<sub>3</sub>), 1742 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta = 4.42$  (t, <sup>3</sup>J = 6 Hz, 8 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 4.35 (t, <sup>3</sup>J = 6 Hz, 16 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 3.43 (t, <sup>3</sup>J = 6 Hz, 8 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.38 (t, <sup>3</sup>J = 6 Hz, 16 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 2.23 (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 1.96 (m, 16 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>) ppm.



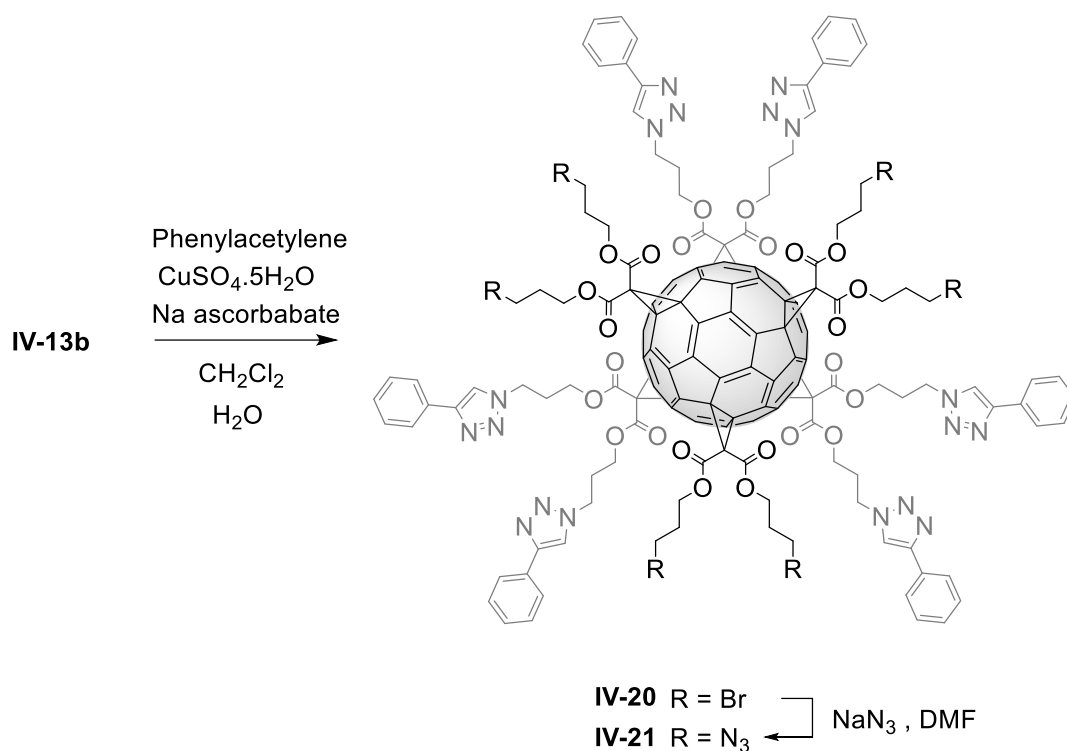
**Compound IV-17:** A solution of **IV-13a** (150 mg, 66  $\mu\text{mol}$ ), phenylacetylene (0.08 mL, 0.76 mmol),  $\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$  (5 mg, 19  $\mu\text{mol}$ ) and sodium ascorbate (10 mg, 50  $\mu\text{mol}$ ) in  $\text{CH}_2\text{Cl}_2/\text{H}_2\text{O}$  (2 : 1 mL) was stirred at 30°C for 15 h, then  $\text{CH}_2\text{Cl}_2$  was added, the organic layer washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 98:2) yielded **IV-17** (179 mg, 62  $\mu\text{mol}$ , 94%) as an orange glassy solid. IR (neat):  $\nu = 1741$  (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 246$  ( $1.5 \times 10^5$ ), 266 (sh,  $8 \times 10^4$ ), 282 ( $7 \times 10^4$ ), 317 ( $4 \times 10^4$ ), 336 ( $3 \times 10^4$ ) nm.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 7.86$  (s, 3 H,  $H_{\text{triazole}}$ ), 7.82 (s, 3 H,  $H_{\text{triazole}}$ ), 7.79 (m, 12 H,  $\text{Ar}H_o$ ), 7.36 (m, 12 H,  $\text{Ar}H_m$ ), 7.30 (m, 6 H,  $\text{Ar}H_p$ ), 4.40 (m, 30 H,  $\text{CH}_2\text{CH}_2\text{OOC}$  and  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 4.28 (q,  $^3J = 7$  Hz, 6 H,  $\text{COOCH}_2\text{CH}_3$ ), 3.42 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ), 2.34 (m, 12 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.21 (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{Br}$ ), 1.28 (t,  $^3J = 7$  Hz, 9 H,  $\text{CH}_2\text{CH}_3$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 163.6$ , 163.5 (two peaks), 163.4, 147.9 (two peaks), 146.2, 146.0 (two peaks), 145.7 (two peaks), 145.4, 141.4 (two peaks), 141.2, 141.1 (two peaks), 141.0, 130.5 (two peaks), 128.8 (two peaks), 128.2, 125.8, 125.7, 120.1 (two peaks), 69.3, 69.2, 69.1, 69.0, 64.6, 63.7, 63.4, 46.8, 45.4, 45.3, 31.3, 29.3, 29.0, 14.1 ppm. MS-MALDI-TOF: 2892.8 ( $[\text{M}+\text{H}]^+$ , calcd for  $\text{C}_{159}\text{H}_{106}\text{O}_{24}\text{N}_{18}\text{Br}_3$ : 2892.5).

**Compound IV-18:** **IV-17** (79 mg, 27  $\mu\text{mol}$ ) and  $\text{NaN}_3$  (10 mg, 150  $\mu\text{mol}$ ) in DMF (2 mL) were stirred at room temperature. After 17 h, the product was precipitated with  $\text{H}_2\text{O}/\text{Et}_2\text{O}$  and resolubilized in  $\text{CH}_2\text{Cl}_2$ , the organic layer washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 98:2) yielded **IV-18** (73 mg, 26  $\mu\text{mol}$ , 96%) as an orange glassy solid. IR (neat):  $\nu = 2099$  ( $\text{N}_3$ ), 1741 (C=O)  $\text{cm}^{-1}$ .  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 7.86$  (s, 3 H,  $H_{\text{triazole}}$ ), 7.83 (s, 3 H,  $H_{\text{triazole}}$ ), 7.81 (d,  $^3J = 8$  Hz, 6 H,  $\text{Ar}H_o$ ), 7.78 (d,  $^3J = 8$  Hz, 6 H,  $\text{Ar}H_o$ ), 7.38 (t,  $^3J = 8$  Hz, 6 H,  $\text{Ar}H_m$ ), 7.36 (t,  $^3J = 8$  Hz, 6 H,  $\text{Ar}H_m$ ), 7.30 (t,  $^3J = 8$  Hz, 6 H,  $\text{Ar}H_p$ ), 4.47 (t,  $^3J = 6$  Hz, 12 H,  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 4.44 (t,  $^3J = 6$  Hz, 12 H,  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 4.36 (m, 18 H,

CH<sub>2</sub>CH<sub>2</sub>OOC), 4.28 (q, <sup>3</sup>J = 7 Hz, 6 H, COOCH<sub>2</sub>CH<sub>3</sub>), 3.37 (t, <sup>3</sup>J = 6 Hz, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 2.37 (m, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 2.32 (m, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 1.94 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 1.28 (t, <sup>3</sup>J = 7 Hz, 9 H, CH<sub>2</sub>CH<sub>3</sub>) ppm.



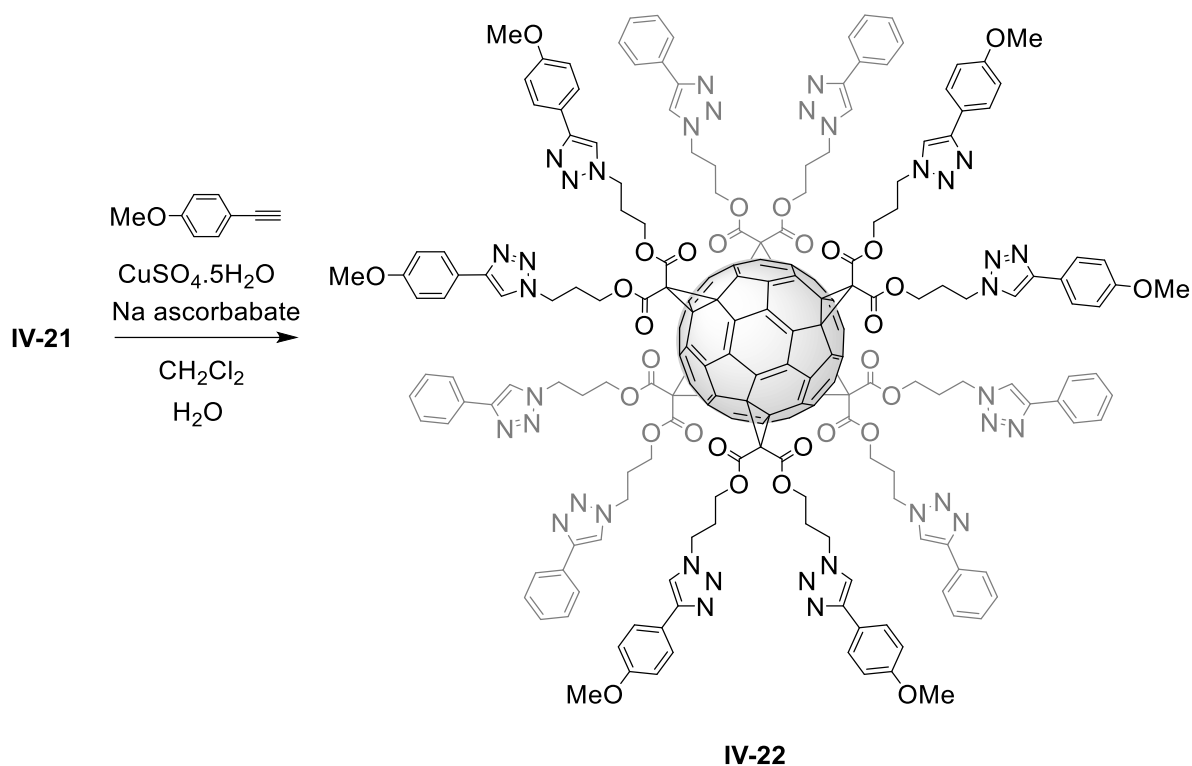
**Compound IV-19:** IV-18 (73 mg, 26 μmol) was added to a solution of 4-ethynylanisole (20 mg, 155 μmol), CuSO<sub>4</sub>·5H<sub>2</sub>O (1 mg, 5 μmol) and sodium ascorbate (3 mg, 15 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (2/1 mL). The resulting mixture was stirred at 30°C for 15 h, then a small amount of reactant and catalyst were added. After 4 h, CH<sub>2</sub>Cl<sub>2</sub> was added to the mixture, the organic layer washed with H<sub>2</sub>O and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 97:3) yielded IV-19 (80 mg, 25 μmol, 92 %) as an orange glassy solid. IR (neat): ν = 1741 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>): λ<sub>max</sub>(ε) = 247 (2x10<sup>5</sup>), 266 (sh, 1x10<sup>5</sup>), 282 (sh, 8x10<sup>4</sup>), 317 (4x10<sup>4</sup>), 336 (3x10<sup>4</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): δ = 7.85 (s, 3 H, H<sub>triazole-Ph</sub>), 7.83 (s, 3 H, H<sub>triazole-Ph</sub>), 7.79 (m, 12 H, ArH<sub>o</sub>), 7.78 (d, <sup>3</sup>J = 8 Hz, 6 H, MeOArH<sub>o</sub>), 7.65 (s, 3 H, H<sub>triazole-PhOMe</sub>), 7.36 (m, 12 H, ArH<sub>m</sub>), 7.29 (m, 6 H, ArH<sub>p</sub>), 6.92 (d, <sup>3</sup>J = 8 Hz, 6 H, MeOArH<sub>m</sub>), 4.45 (m, 12 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 4.37 (m, 18 H, COOCH<sub>2</sub>CH<sub>2</sub> and CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 4.29 (m, 12 H, COOCH<sub>2</sub>), 3.80 (s, 9 H, ArOCH<sub>3</sub>), 2.34 (m, 12 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 2.27 (m, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.25 (t, <sup>3</sup>J = 7 Hz, CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz): δ = 163.5 (two peaks), 163.4 (two peaks), 159.6, 147.9 (two peaks), 147.7, 146.3 (two peaks), 146.0, 145.9, 145.8 (two peaks), 145.5, 145.4, 141.6, 141.5, 141.2, 141.1 (two peaks), 141.0, 134.5, 130.5 (two peaks), 128.8, 128.2, 127.0, 125.7 (two peaks), 124.3, 123.2, 122.0, 120.1, 119.2, 114.3, 69.3, 69.2, 69.1 (two peaks), 63.8, 63.7, 63.5, 63.4, 55.3, 46.8, 46.7, 45.5, 45.4, 29.3, 14.1 ppm. MS-MALDI-TOF: 3176.3 ([M+2H]<sup>+</sup>, calcd for C<sub>186</sub>H<sub>131</sub>O<sub>27</sub>N<sub>27</sub>: 3176.0). Anal. C<sub>214</sub>H<sub>152</sub>O<sub>28</sub>N<sub>36</sub>•1/2CHCl<sub>3</sub>•1/2CH<sub>2</sub>Cl<sub>2</sub> (3276.34): calcd. C 68.55, H 4.01, N 11.54 ; found C 68.66, H 3.62, N 11.54.



**Compound IV-20:** A solution of **IV-13b** (94 mg, 37  $\mu\text{mol}$ ), phenylacetylene (0.05 mL, 0.47 mmol),  $\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$  (3 mg, 11  $\mu\text{mol}$ ) and sodium ascorbate (4 mg, 22  $\mu\text{mol}$ ) in  $\text{CH}_2\text{Cl}_2/\text{H}_2\text{O}$  (2/1 mL) was stirred at 30°C for 15 h, then  $\text{CH}_2\text{Cl}_2$  was added, the organic layer washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 98:2) yielded **IV-20** (116 mg, 36  $\mu\text{mol}$ , 99%) as an orange glassy solid. IR (neat):  $\nu = 1743$  (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 246$  ( $2.5 \times 10^5$ ), 267 (sh,  $1.3 \times 10^5$ ), 281 (sh,  $1.1 \times 10^5$ ), 318 (sh,  $6 \times 10^4$ ), 336 (sh,  $5 \times 10^4$ ) nm.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 7.86$  (s, 3 H,  $H_{\text{triazole-Ph}}$ ), 7.83 (s, 3 H,  $H_{\text{triazole-Ph}}$ ), 7.79 (m, 12 H,  $\text{ArH}_o$ ), 7.37 (m, 12 H,  $\text{ArH}_m$ ), 7.30 (m, 6 H,  $\text{ArH}_p$ ), 4.49-4.34 (m, 36 H,  $\text{CH}_2\text{CH}_2\text{OOC}$  &  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 3.41 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ), 3.39 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ), 2.38 (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.32 (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.22 (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{Br}$ ), 2.19 (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{Br}$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 163.5$ , 163.4 (three peaks), 147.9 (two peaks), 145.9 (two peaks), 145.8 (two peaks), 145.7, 141.3, 141.2 (three peaks), 141.1, 141.0, 130.5 (two peaks), 128.8, 128.2, 125.7 (two peaks), 120.1 (two peaks), 69.2, 69.1 (three peaks), 64.8, 63.7 (two peaks), 46.8, 45.4, 45.3, 31.2 (two peaks), 29.3, 29.1 (two peaks) ppm. MS-MALDI-TOF: 3171.1 ( $[\text{M}+\text{H}]^+$ , calcd for  $\text{C}_{162}\text{H}_{109}\text{O}_{24}\text{N}_{18}\text{Br}_6$ : 3171.3).

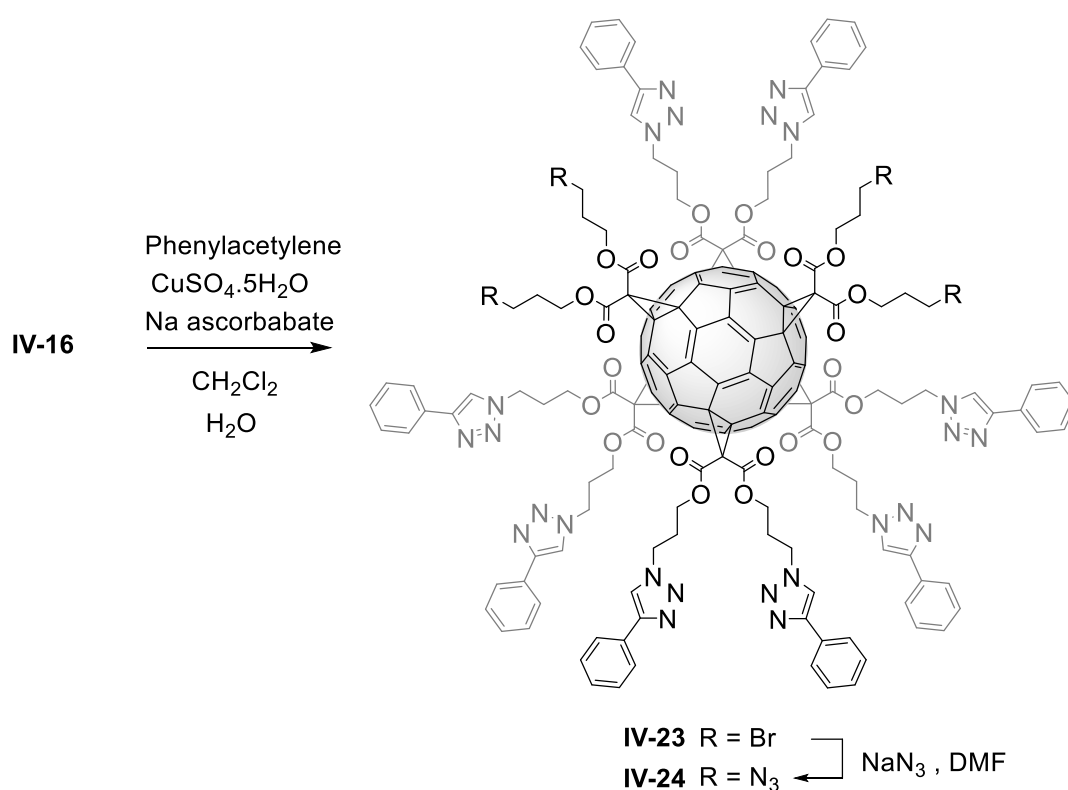
**Compound IV-21:** A solution of **IV-20** (113 mg, 35  $\mu\text{mol}$ ) and  $\text{NaN}_3$  (23 mg, 356  $\mu\text{mol}$ ) in DMF (2 mL) was stirred at room temperature. After 17 h, the product was precipitated with  $\text{H}_2\text{O}/\text{Et}_2\text{O}$  and resolubilized in  $\text{CH}_2\text{Cl}_2$ , the organic layer washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 98:2) yielded **IV-21** (103 mg, 35  $\mu\text{mol}$ , 98%) as an orange glassy solid. IR (neat):  $\nu = 2098$  ( $\text{N}_3$ ), 1742 (C=O)  $\text{cm}^{-1}$ .  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 7.86$  (s, 3 H,  $H_{\text{triazole}}$ ), 7.83 (s, 3 H,  $H_{\text{triazole}}$ ), 7.79 (m, 12 H,  $\text{ArH}_o$ ), 7.35 (m, 12 H,

ArH<sub>m</sub>), 7.30 (m, 6 H, ArH<sub>p</sub>), 4.45 (m, 12 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 4.34 (m, 24 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 3.36 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 3.33 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 2.34 (m, 12 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 1.94 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 1.90 (quint., <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>) ppm.



**Compound IV-22:** **IV-21** (103 mg, 35  $\mu$ mol) was added to a solution of 4-ethynylanisole (57 mg, 428  $\mu$ mol), CuSO<sub>4</sub>.5H<sub>2</sub>O (3 mg, 11  $\mu$ mol) and sodium ascorbate (4 mg, 22  $\mu$ mol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (2/1 mL). The resulting mixture was stirred at 30°C for 15 h, then CH<sub>2</sub>Cl<sub>2</sub> was added to the mixture, the organic layer washed with H<sub>2</sub>O and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 97:3) yielded **IV-22** (117 mg, 31  $\mu$ mol, 88%) as an orange glassy solid. IR (neat):  $\nu$  = 1742 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon)$  = 248 (2.5x10<sup>5</sup>), 318 (sh, 5x10<sup>4</sup>), 335 (sh, 4x10<sup>4</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  = 7.82 (s, 6 H, H<sub>triazole-Ph</sub>), 7.76 (d, <sup>3</sup>J = 7 Hz, 12 H, ArH<sub>o</sub>), 7.71 (s, 6 H, H<sub>triazole-PhOMe</sub>), 7.67 (d, <sup>3</sup>J = 9 Hz, 12 H, MeOArH<sub>o</sub>), 7.34 (t, <sup>3</sup>J = 7 Hz, 12 H, ArH<sub>m</sub>), 7.28 (d, <sup>3</sup>J = 7 Hz, 6 H, ArH<sub>p</sub>), 6.87 (d, <sup>3</sup>J = 9 Hz, 12 H, MeOArH<sub>m</sub>), 4.41 (t, <sup>3</sup>J = 7 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 4.39 (t, <sup>3</sup>J = 7 Hz, 12 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 4.33 (t, <sup>3</sup>J = 6 Hz, 24 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 3.77 (s, 18 H, ArOCH<sub>3</sub>), 2.29 (m, 24 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  = 163.4, 159.6, 147.9, 147.8, 145.9, 141.3, 130.5, 128.9, 128.2, 127.0, 125.7, 123.2, 120.1, 119.3, 114.3, 69.2, 63.8, 55.3, 46.8 (two peaks), 45.5, 29.3 ppm. MS-MALDI-TOF: 3735.8 ([M+H]<sup>+</sup>, calcd for C<sub>216</sub>H<sub>156</sub>O<sub>30</sub>N<sub>36</sub>: 3736.2). Anal. C<sub>216</sub>H<sub>156</sub>O<sub>30</sub>N<sub>36</sub>•0.8CHCl<sub>3</sub> (3831.28): calcd. C 67.96 H 4.12 N 13.16 ; found C 68.00 H 4.00 N 13.07.

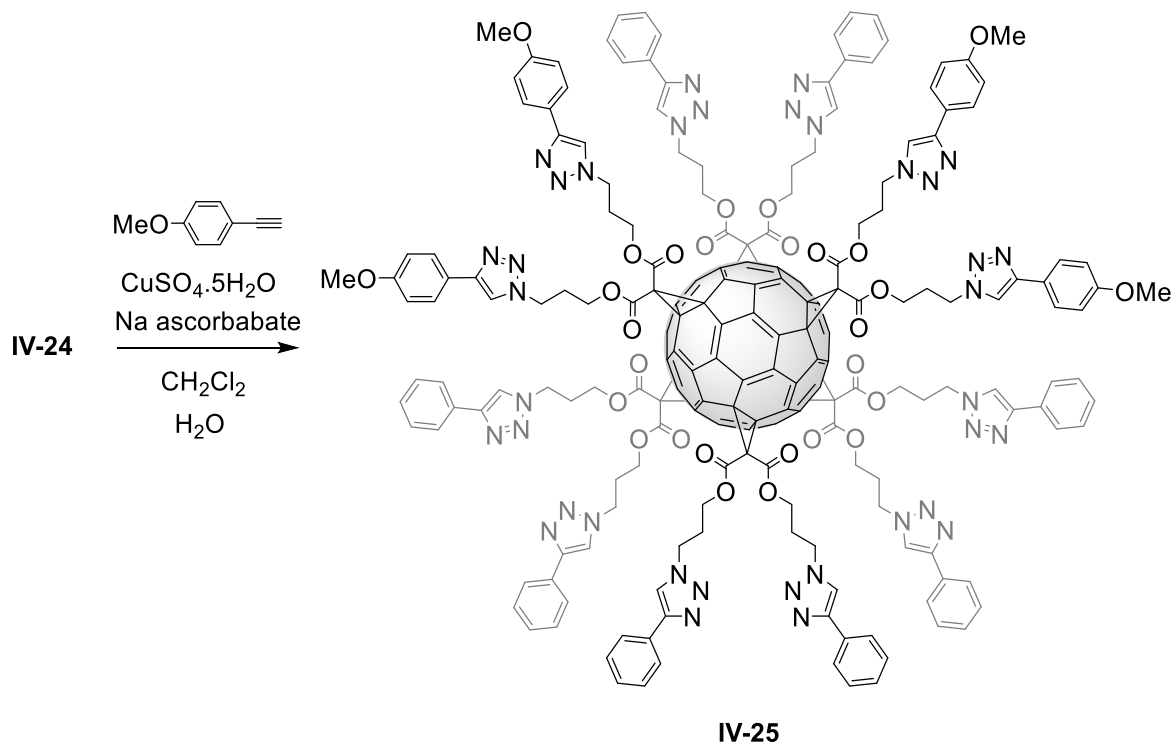




**Compound IV-23:** A solution of **IV-16** (265 mg, 107  $\mu\text{mol}$ ), phenylacetylene (0.19 mL, 1.7 mmol),  $\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$  (10 mg, 42  $\mu\text{mol}$ ) and sodium ascorbate (17 mg, 85  $\mu\text{mol}$ ) in  $\text{CH}_2\text{Cl}_2/\text{H}_2\text{O}$  (2/1 mL) was stirred at 30°C for 15 h, then  $\text{CH}_2\text{Cl}_2$  was added, washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 98:2) yielded **IV-23** (322 mg, 98  $\mu\text{mol}$ , 91%) as an orange glassy solid. IR (neat):  $\nu = 1742$  (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 246$  ( $2.4 \times 10^5$ ), 284 (sh,  $9 \times 10^4$ ), 322 (sh,  $5 \times 10^4$ ), 338 (sh,  $4 \times 10^4$ ) nm.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 7.79$  (m, 24 H), 7.34 (m, 24 H), 4.44 (m, 16 H,  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 4.36 (m, 24 H,  $\text{CH}_2\text{CH}_2\text{OOC}$ ), 3.39 (m, 8 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ), 2.23 (m, 16 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.18 (m, 8 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{Br}$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 163.5$  (two peaks), 163.4 (two peaks), 163.3, 147.9 (three peaks), 145.9 (three peaks), 145.8 (two peaks), 145.7, 141.3 (two peaks), 141.2 (three peaks), 141.1 (three peaks), 130.5 (two peaks), 128.8, 128.2, 125.7 (two peaks), 120.1 (two peaks), 69.2, 69.1 (two peaks), 64.8, 63.7 (two peaks), 46.8, 45.4 (three peaks), 45.3, 31.2 (two peaks), 29.3, 29.1 ppm. MS-MALDI-TOF: 3300.1 ( $[\text{M}+\text{H}]^+$ , calcd for  $\text{C}_{178}\text{H}_{121}\text{O}_{24}\text{N}_{24}\text{Br}_4$ : 3299.6).

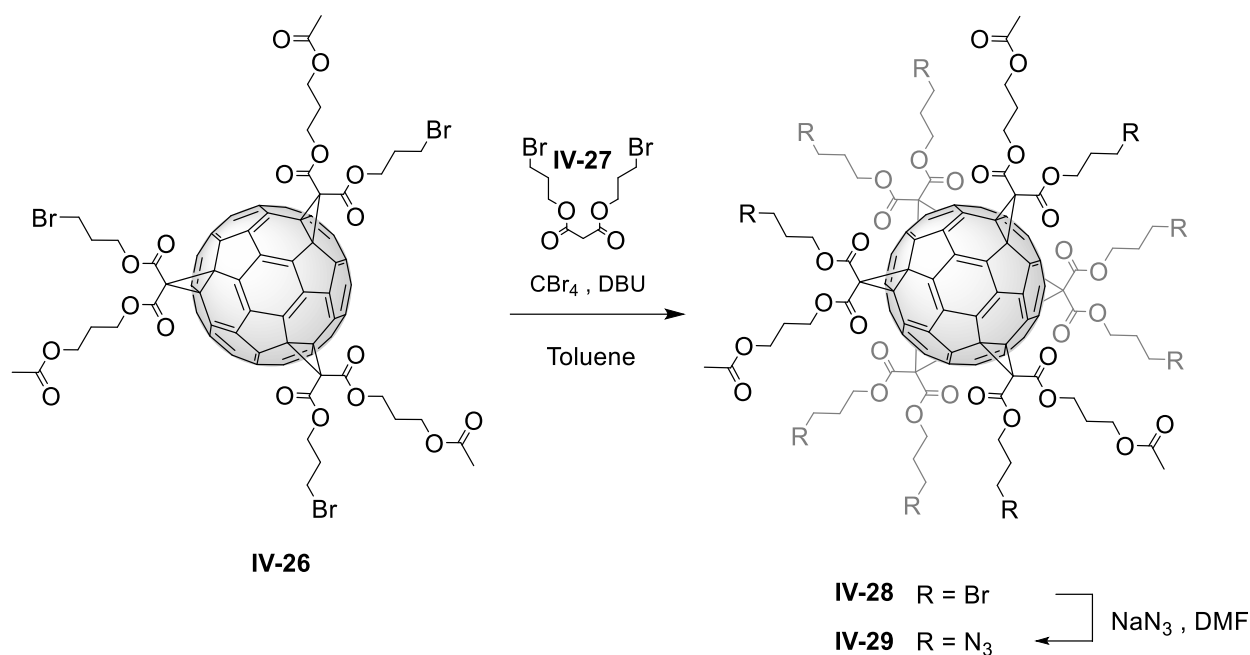
**Compound IV-24:** A solution of **IV-23** (149 mg, 45  $\mu\text{mol}$ ) and  $\text{NaN}_3$  (18 mg, 271  $\mu\text{mol}$ ) in DMF (2 mL) was stirred at room temperature. After 17 h, the product was precipitated with  $\text{H}_2\text{O}/\text{Et}_2\text{O}$  and resolubilized in  $\text{CH}_2\text{Cl}_2$ , washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 98:2) yielded **IV-24** (139 mg, 44  $\mu\text{mol}$ , 98%) as an orange glassy solid. IR (neat):  $\nu = 2098$  ( $\text{N}_3$ ), 1741 (C=O)  $\text{cm}^{-1}$ .  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 7.86\text{--}7.81$  (4s, 8 H,  $H_{\text{triazole}}$ ), 7.75 (m, 16 H,  $\text{ArH}_o$ ), 7.30 (m, 24 H,  $\text{ArH}_{m,p}$ ), 4.41 (m, 16 H,

CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 4.31 (m, 24 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 3.30 (t, <sup>3</sup>J = 7 Hz, 8 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 2.30 (m, 16 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 1.87 (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>) ppm.



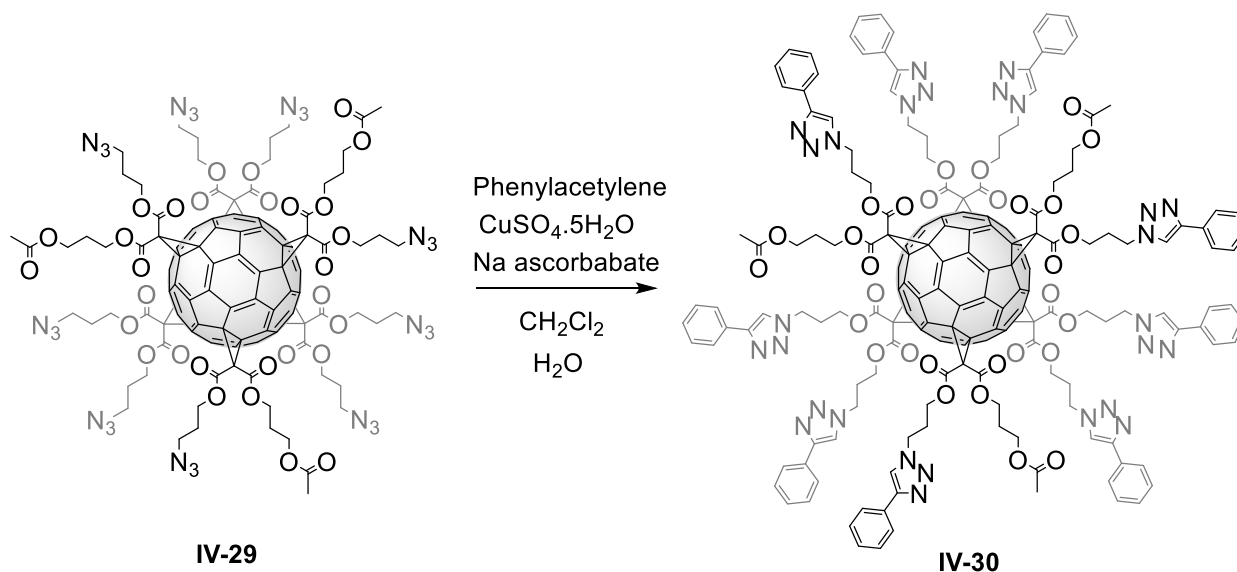
**Compound IV-25:** IV-24 (139 mg, 44 μmol) was added to a solution of 4-ethynylanisole (0.06 mL, 0.5 mmol), CuSO<sub>4</sub>·5H<sub>2</sub>O (2 mg, 9 μmol) and sodium ascorbate (3 mg, 18 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (2/1 mL). The resulting mixture was stirred at 30°C for 24 h, then a small amount of reactant and catalyst were added. After 18 h, CH<sub>2</sub>Cl<sub>2</sub> was added to the mixture, the organic layer washed with H<sub>2</sub>O and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 97:3) yielded IV-25 (122 mg, 33 μmol, 75%) as an orange glassy solid. IR (neat):  $\nu = 1742$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 247$  (2.2x10<sup>5</sup>), 284 (sh, 7x10<sup>4</sup>), 321 (sh, 4x10<sup>4</sup>), 336 (sh, 3x10<sup>4</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.82$  (s, 8 H, *H*<sub>Triazole Ph</sub>), 7.76 (d, <sup>3</sup>J = 7 Hz, 16 H, *ArH<sub>o</sub>*), 7.72 (s, 4 H, *H*<sub>Triazole PhOMe</sub>), 7.67 (d, <sup>3</sup>J = 9 Hz, 8 H, H<sub>3</sub>CO*ArH<sub>o</sub>*), 7.34 (t, <sup>3</sup>J = 7 Hz, 8 H, *ArH<sub>m</sub>*), 7.28 (d, <sup>3</sup>J = 7 Hz, 16 H, *ArH<sub>p</sub>*), 6.87 (d, <sup>3</sup>J = 9 Hz, 8 H, H<sub>3</sub>CO*ArH<sub>m</sub>*), 4.41 (t, <sup>3</sup>J = 7 Hz, 16 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 4.39 (t, <sup>3</sup>J = 7 Hz, 8 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 4.33 (t, <sup>3</sup>J = 6 Hz, 24 H, CH<sub>2</sub>CH<sub>2</sub>OOC), 3.77 (s, 12 H, OCH<sub>3</sub>), 2.29 (m, 24 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.4, 159.6, 147.9, 147.8, 145.9, 141.3, 130.5, 128.9, 128.2, 127.0, 125.7, 123.2, 120.1, 119.3, 114.3, 69.2, 63.8, 55.3, 46.8$  (two peaks), 45.5, 29.3 ppm. MS-MALDI-TOF : 3676.7 ([M+H]<sup>+</sup>, calcd for C<sub>214</sub>H<sub>153</sub>N<sub>36</sub>O<sub>28</sub>: 3676.2). Anal. C<sub>214</sub>H<sub>152</sub>O<sub>28</sub>N<sub>36</sub>•CHCl<sub>3</sub> (3795.11) : calcd. C 68.04, H 4.06, N 13.29 ; found C 68.05, H 3.87, N 13.38.



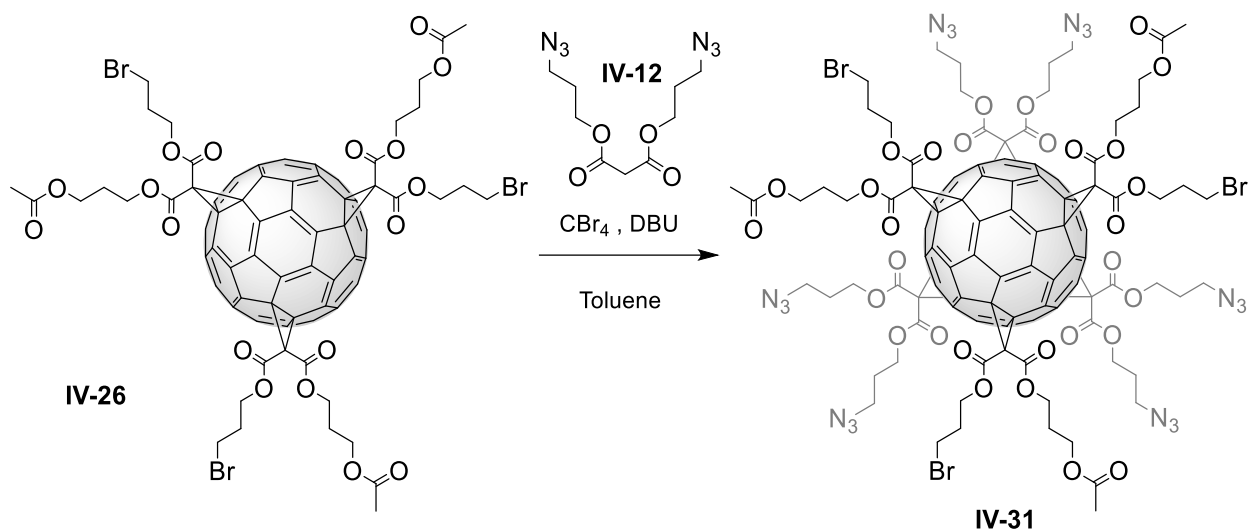


**Compound IV-28:** DBU (0.11 mL, 0.76 mmol) was added to a solution of **IV-26** (80 mg, 47  $\mu\text{mol}$ ), **IV-27** (98 mg, 284  $\mu\text{mol}$ ) and CBr<sub>4</sub> (784 mg, 2.4 mmol) in toluene (25 mL). The resulting mixture was stirred at room temperature for 2 h, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/Et<sub>2</sub>O, 95:5) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/Et<sub>2</sub>O, 98:2) yielded **IV-28** (86 mg, 31.6  $\mu\text{mol}$ , 67%) as an orange glassy solid. IR (neat):  $\nu = 1735$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 245$  (1.2x10<sup>5</sup>), 270 (sh, 8.6x10<sup>4</sup>), 281 (9.0x10<sup>4</sup>), 317 (sh, 5.6x10<sup>4</sup>), 336 (sh, 4.6x10<sup>4</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.43$  (t, <sup>3</sup>J = 6 Hz, 18 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 4.37 (t, <sup>3</sup>J = 6 Hz, 6 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OOCCH<sub>3</sub>), 4.14 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOCCH<sub>3</sub>), 3.44 (m, 18 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.25 (m, 18 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 2.06 (m, 15 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OOCCH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 170.9$ , 163.5, 146.0, 145.9, 145.8, 145.7 (two peaks), 145.6, 141.2, 141.1 (two peaks), 140.0, 69.1 (two peaks), 64.7, 63.8, 60.6, 45.2, 31.3, 29.1, 27.8, 20.9 ppm. MALDI-TOF-MS: 2721.5 (100%, [M]<sup>+</sup>, calcd for C<sub>120</sub>H<sub>81</sub>O<sub>30</sub>Br<sub>9</sub>: 2721.7), 2585.6 (39%, [M+H-(OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br)]<sup>+</sup>, calcd for C<sub>117</sub>H<sub>76</sub>O<sub>29</sub>Br<sub>8</sub>: 2585.8), 2377.7 (28%, [M-(malonate-Br)]<sup>+</sup>, calcd for C<sub>111</sub>H<sub>69</sub>O<sub>26</sub>Br<sub>7</sub>: 2377.8).

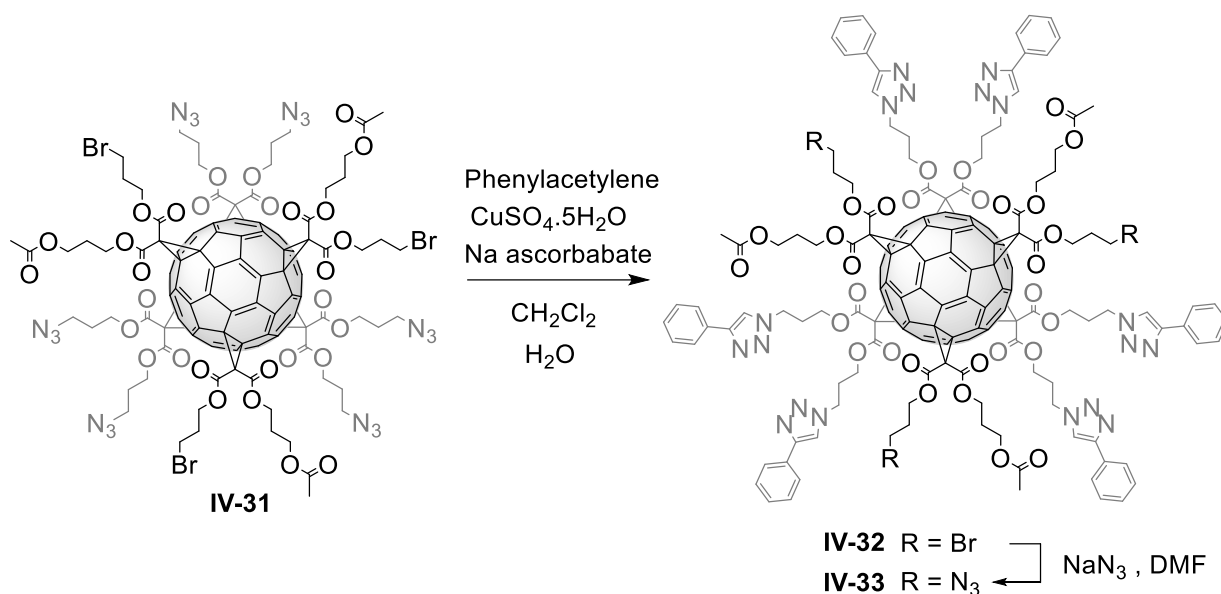
**Compound IV-29:** A solution of **IV-28** (66 mg, 24.2  $\mu\text{mol}$ ) and NaN<sub>3</sub> (28 mg, 436  $\mu\text{mol}$ ) in DMF (2 mL) was stirred at room temperature for 16 h. Then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/Et<sub>2</sub>O, 92:8) and concentrated. Gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) yielded **IV-29** (56 mg, 23.5  $\mu\text{mol}$ , 97%) as an orange glassy solid. IR (neat):  $\nu = 2097$  (N<sub>3</sub>), 1740 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 300 MHz):  $\delta = 4.37$  (t, <sup>3</sup>J = 6 Hz, 24 H, COOCH<sub>2</sub>CH<sub>2</sub>), 4.14 (t, <sup>3</sup>J = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOCCH<sub>3</sub>), 3.39 (m, 18 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.07 (m, 15 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OOCCH<sub>3</sub>), 1.97 (quint., <sup>3</sup>J = 6 Hz, 18 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>) ppm.



**Compound IV-30:** A solution of **IV-29** (55 mg, 23.1  $\mu\text{mol}$ ), phenylacetylene (0.04 mL, 323  $\mu\text{mol}$ ), CuSO<sub>4</sub>·5H<sub>2</sub>O (3 mg, 11  $\mu\text{mol}$ ) and sodium ascorbate (4 mg, 22  $\mu\text{mol}$ ) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (3/1 mL) was stirred at 30°C for 4 days, then CH<sub>2</sub>Cl<sub>2</sub> was added, the organic layer washed with H<sub>2</sub>O and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>  $\rightarrow$  CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 96:4) yielded **IV-30** (72 mg, 21.8  $\mu\text{mol}$ , 94%) as an orange glassy solid. IR (neat):  $\nu = 1740$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 246$  (2.2x10<sup>5</sup>), 263 (sh, 1.3x10<sup>5</sup>), 282 (sh, 8x10<sup>4</sup>), 318 (sh, 5x10<sup>4</sup>), 337 (sh, 4x10<sup>4</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.82$  (s, 3 H, *H*<sub>triazole</sub>), 7.80-7.75 (m, 24 H, *H*<sub>triazole</sub> & Ar*H*<sub>o</sub>), 7.36 (m, 18 H, Ar*H*<sub>m</sub>), 7.29 (m, 9 H, Ar*H*<sub>p</sub>), 4.44-4.30 (m, 42 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub> & COOCH<sub>2</sub>CH<sub>2</sub>), 4.10 (t, <sup>3</sup>*J* = 6 Hz, 6 H, CH<sub>2</sub>CH<sub>2</sub>OOCCH<sub>3</sub>), 2.29 (m, 18 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 2.01 (m, 15 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OOCCH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 170.9, 163.5, 163.4$  (three peaks), 147.9, 147.8, 146.3 (two peaks), 145.9, 145.8, 145.5, 145.4, 141.6, 141.5, 141.3, 141.2, 141.1 (two peaks), 141.0, 140.9, 130.5 (two peaks), 130.4, 128.8, 128.2, 125.7, 120.0 (two peaks), 69.2 (two peaks), 69.1, 64.0, 63.7 (br), 60.6, 46.9, 46.8, 45.4 (two peaks), 29.4, 29.3, 27.7, 20.9 ppm. MALDI-TOF-MS: 3301.2 ([M+H]<sup>+</sup>, calcd for C<sub>192</sub>H<sub>136</sub>O<sub>30</sub>N<sub>27</sub>: 3301.0). Anal. C<sub>192</sub>H<sub>135</sub>O<sub>30</sub>N<sub>27</sub>·CH<sub>2</sub>Cl<sub>2</sub> (3385.23): calcd. C 68.48 H 4.08 N 11.17; found C 68.66 H 4.03 N 11.07.



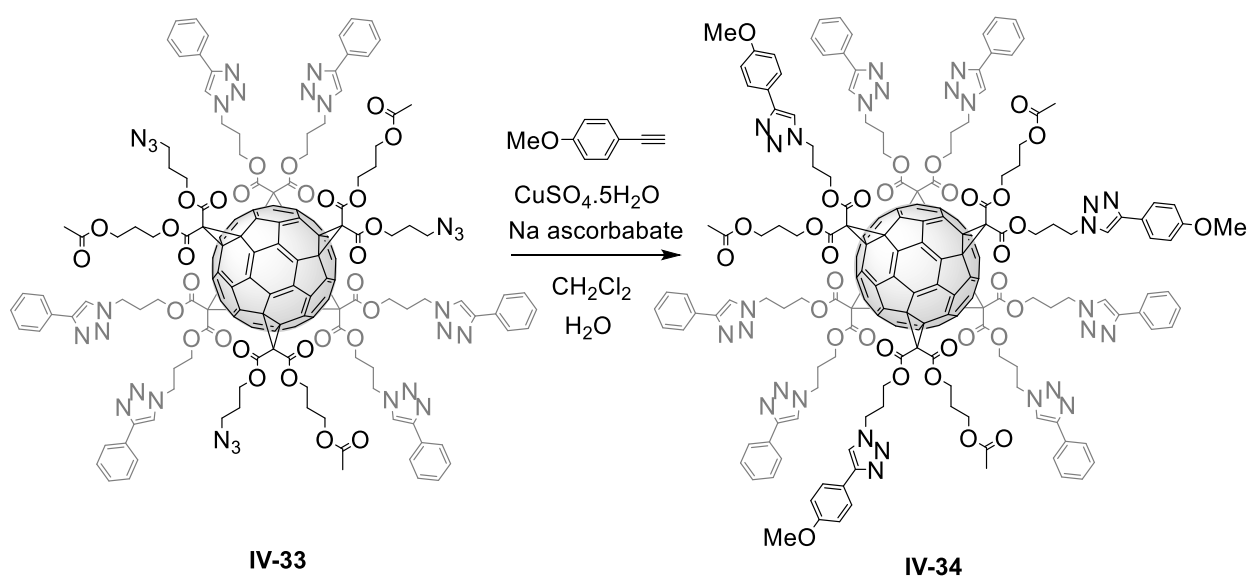
**Compound IV-31:** DBU (0.08 mL, 0.5 mmol) was added to a solution of **IV-26** (55 mg, 32.5  $\mu\text{mol}$ ), **IV-3** (53 mg, 195  $\mu\text{mol}$ ) and  $\text{CBr}_4$  (540 mg, 1.6 mmol) in toluene (15 mL). The resulting mixture was stirred at room temperature for 4 h, then filtered on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{Et}_2\text{O}$ , 9:1) and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{Et}_2\text{O}$ , 95:5) yielded **IV-31** (59 mg, 23.6  $\mu\text{mol}$ , 73%) as an orange glassy solid. IR (neat):  $\nu = 2098$  ( $\text{N}_3$ ),  $1739$  ( $\text{C}=\text{O}$ )  $\text{cm}^{-1}$ .  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 4.43$  (t,  $^3J = 6$  Hz, 6 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{Br}$ ),  $4.38$  (t,  $^3J = 6$  Hz, 18 H,  $\text{COOCH}_2\text{CH}_2$ ),  $4.15$  (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{OOCCH}_3$ ),  $3.43$  (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ),  $3.40$  (t,  $^3J = 6$  Hz, 12 H,  $\text{CH}_2\text{CH}_2\text{N}_3$ ),  $2.25$  (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{Br}$ ),  $2.07$  (m, 15 H, ),  $1.97$  (quint.,  $^3J = 6$  Hz, 12 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_3$ ) ppm.



**Compound IV-32:** A solution of **IV-31** (59 mg, 23.6  $\mu\text{mol}$ ), phenylacetylene (0.03 mL, 238  $\mu\text{mol}$ ),  $\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$  (3 mg, 11  $\mu\text{mol}$ ) and sodium ascorbate (4 mg, 22  $\mu\text{mol}$ ) in  $\text{CH}_2\text{Cl}_2/\text{H}_2\text{O}$  (2 : 1 mL) was stirred at  $30^\circ\text{C}$  for 16 h, then  $\text{CH}_2\text{Cl}_2$  was added, the organic layer washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 98:2) yielded **IV-**

**32** (73 mg, 23.5  $\mu\text{mol}$ , 99%) as an orange glassy solid. IR (neat):  $\nu = 1740$  (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 246$  ( $1.4 \times 10^5$ ), 266 (sh,  $8 \times 10^4$ ), 282 (sh,  $6 \times 10^4$ ), 319 (sh,  $4 \times 10^4$ ), 336 (sh,  $3 \times 10^4$ ) nm.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 7.87$  (s, 3 H,  $H_{\text{triazole}}$ ), 7.83 (s, 3 H,  $H_{\text{triazole}}$ ), 7.81 (d,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_o$ ), 7.78 (d,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_o$ ), 7.38 (t,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_m$ ), 7.36 (t,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_m$ ), 7.30 (t,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_p$ ), 4.48 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 4.44 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 4.37 (m, 24 H,  $\text{COOCH}_2\text{CH}_2$ ), 4.12 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{OOCCH}_3$ ), 3.38 (t,  $^3J = 6$  Hz,  $\text{CH}_2\text{CH}_2\text{Br}$ ), 2.38 (m, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.32 (m, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.18 (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{Br}$ ), 2.06 (m, 15 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{OOCCH}_3$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 170.9$ , 163.5, 163.5, 163.4, 147.9 (two peaks), 146.0 (two peaks), 145.9, 145.8 (four peaks), 145.7, 145.6, 141.4 (two peaks), 141.2, 141.1, 141.0 (two peaks), 130.5, 130.4, 128.8, 128.2 (two peaks), 125.7 (two peaks), 120.1 (two peaks), 69.2, 69.1, 69.0, 64.8, 63.9, 63.7 (br), 60.6, 46.8 (br), 45.3 (two peaks), 31.2, 29.3, 29.1, 27.7, 20.9 ppm. MALDI-TOF-MS: 3108.6 ( $[\text{M}]^+$ , calcd for  $\text{C}_{168}\text{H}_{118}\text{O}_{30}\text{N}_{18}\text{Br}_3$ : 3108.6).

**Compound IV-33:** A solution of **IV-32** (72 mg, 23.2  $\mu\text{mol}$ ) and  $\text{NaN}_3$  (14 mg, 208  $\mu\text{mol}$ ) in DMF (2 mL) was stirred at room temperature. After 16 h, the product was precipitated with  $\text{H}_2\text{O}/\text{Et}_2\text{O}$  and resolubilized in  $\text{CH}_2\text{Cl}_2$ , the organic layer washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 96:4) followed by gel permeation chromatography (Biobeads SX-1,  $\text{CH}_2\text{Cl}_2$ ) yielded **IV-33** (65 mg, 21.7  $\mu\text{mol}$ , 94%) as an orange glassy solid. IR (neat):  $\nu = 2098$  ( $\text{N}_3$ ), 1736 (C=O)  $\text{cm}^{-1}$ .  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 300 MHz):  $\delta = 7.86$  (s, 3 H,  $H_{\text{triazole}}$ ), 7.83 (s, 3 H,  $H_{\text{triazole}}$ ), 7.79 (m, 12 H,  $\text{ArH}_o$ ), 7.37 (m, 12 H,  $\text{ArH}_m$ ), 7.30 (m, 6 H,  $\text{ArH}_p$ ), 4.48 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 4.44 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 4.37 (m, 24 H,  $\text{COOCH}_2\text{CH}_2$ ), 4.12 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{OOCCH}_3$ ), 3.33 (t,  $^3J = 6$  Hz,  $\text{CH}_2\text{CH}_2\text{N}_3$ ), 2.38 (m, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.32 (m, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.05 (m, 15 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{OOCCH}_3$ ), 1.91 (quint.,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_3$ ), ppm.



**Compound IV-34:** A solution of **IV-33** (65 mg, 21.7  $\mu\text{mol}$ ), 4-ethynylanisole (26 mg, 195  $\mu\text{mol}$ ),  $\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$  (3 mg, 11  $\mu\text{mol}$ ) and sodium ascorbate (4 mg, 22  $\mu\text{mol}$ ) in  $\text{CH}_2\text{Cl}_2/\text{H}_2\text{O}$  (3/1 mL) was stirred at 30°C for 3.5 days, then  $\text{CH}_2\text{Cl}_2$  was added, the organic layer washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 96:4) yielded **IV-34** (66 mg, 19.5  $\mu\text{mol}$ , 90%) as an orange glassy solid. IR (neat):  $\nu = 1738$  (C=O)  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 247$  ( $1.9 \times 10^5$ ), 265 (sh,  $1.2 \times 10^5$ ), 282 (sh,  $7 \times 10^4$ ), 319 (sh,  $4 \times 10^4$ ), 337 (sh,  $3 \times 10^4$ ) nm.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 7.82$  (s, 3 H,  $H_{\text{triazole-Ph}}$ ), 7.80 (s, 3 H,  $H_{\text{triazole-Ph}}$ ), 7.77 (d,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_o$ ), 7.75 (d,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_o$ ), 7.71 (d,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_o\text{OMe}$ ), 7.70 (s, 3 H,  $H_{\text{triazole-PhOMe}}$ ), 7.36 (t,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_m$ ), 7.34 (t,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_m$ ), 7.29 (m, 6 H,  $\text{ArH}_p$ ), 6.91 (d,  $^3J = 8$  Hz, 6 H,  $\text{ArH}_m\text{OMe}$ ), 4.42-4.30 (m, 42 H,  $\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$  &  $\text{COOCH}_2\text{CH}_2$ ), 4.10 (t,  $^3J = 6$  Hz, 6 H,  $\text{CH}_2\text{CH}_2\text{OOCCH}_3$ ), 3.79 (s, 9 H,  $\text{ArOCH}_3$ ), 2.30 (m, 18 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.02 (m, 15 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{OOCCH}_3$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 170.9$ , 163.5, 163.4 (three peaks), 159.6, 147.9, 147.7, 146.3 (two peaks), 145.9, 145.8, 145.5, 145.4, 141.6, 141.5, 141.3, 141.2, 141.1 (two peaks), 141.0, 140.9, 130.5, 130.4, 128.8, 128.2, 127.0, 125.7 (two peaks), 125.6, 123.2, 120.0 (two peaks), 119.2, 114.3, 69.2 (two peaks), 69.1, 64.0, 63.7 (two peaks), 60.6, 55.3, 46.8, 45.4 (two peaks), 29.4, 29.3, 27.7, 20.9 ppm. MALDI-TOF-MS: 3391.1 ( $[\text{M}+\text{H}]^+$ , calcd for  $\text{C}_{195}\text{H}_{142}\text{O}_{33}\text{N}_{27}$ : 3391.0). Anal.  $\text{C}_{195}\text{H}_{141}\text{O}_{33}\text{N}_{27} \cdot \text{CH}_2\text{Cl}_2$  (3475.31): calcd. C 67.74 H 4.15 N 11.88; found C 67.77 H 4.15 N 10.78.



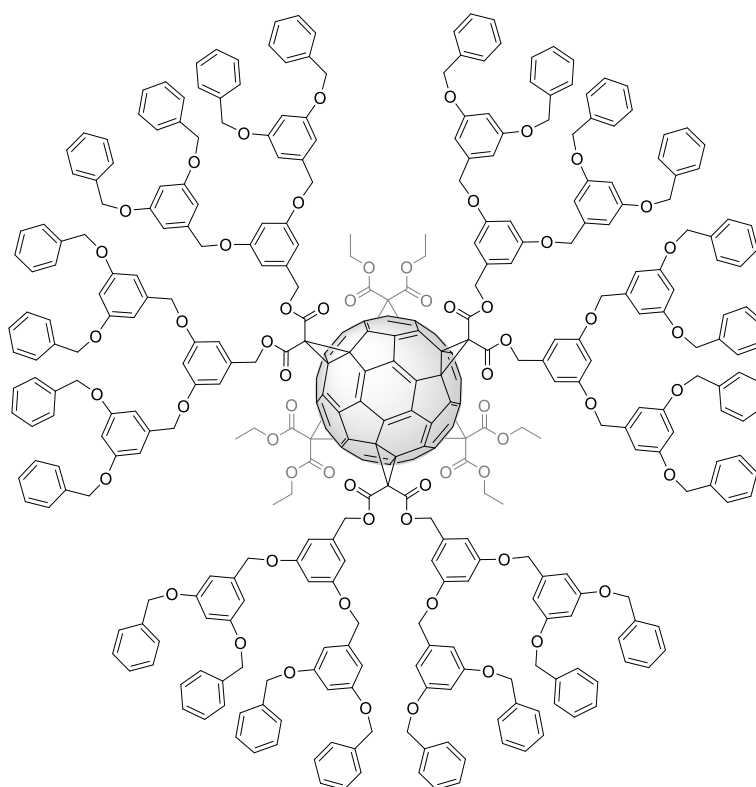


# **Chapter V : Elaboration of dendrimers based on C<sub>60</sub> hexa-adducts.**

## 1. Introduction.

Dendrimers are remarkable polyfunctional molecular structures. Due to their branched architecture, dendrimers are able to generate specific properties.<sup>[1,2]</sup> Furthermore, the high number of functional groups at the periphery of the dendrimer is very attractive for the tuning of potential properties.

Since the first reported C<sub>60</sub> derivative functionalized with a dendritic macromolecule,<sup>[3]</sup> the field of fullerodendrimers has developed concurrently with the fullerene chemistry allowing the formation of more and more elaborated dendritic structures.<sup>[4-7]</sup> The spherical shape of C<sub>60</sub> has proven to be an efficient scaffold for the elaboration of globular dendrimers even with low generation dendrons. *T<sub>h</sub>*-symmetrical hexa-adducts of C<sub>60</sub> are particularly interesting in this field with their multiplicity of 12 and their 3D architectures. Formation of *T<sub>h</sub>*-hexa-adducts of C<sub>60</sub> are sensitive to steric hindrances and when malonates are bearing bulky substituents yields are low or the hexa-adducts are not formed. Only few examples of C<sub>60</sub> hexa-adducts dendrimers have been reported in the literature, among them C<sub>60</sub> hexa-adducts bearing two different type of addends (Figure V-1).<sup>[5]</sup> In this last case, the regiochemistry of multiple additions on C<sub>60</sub> is an additional problem.

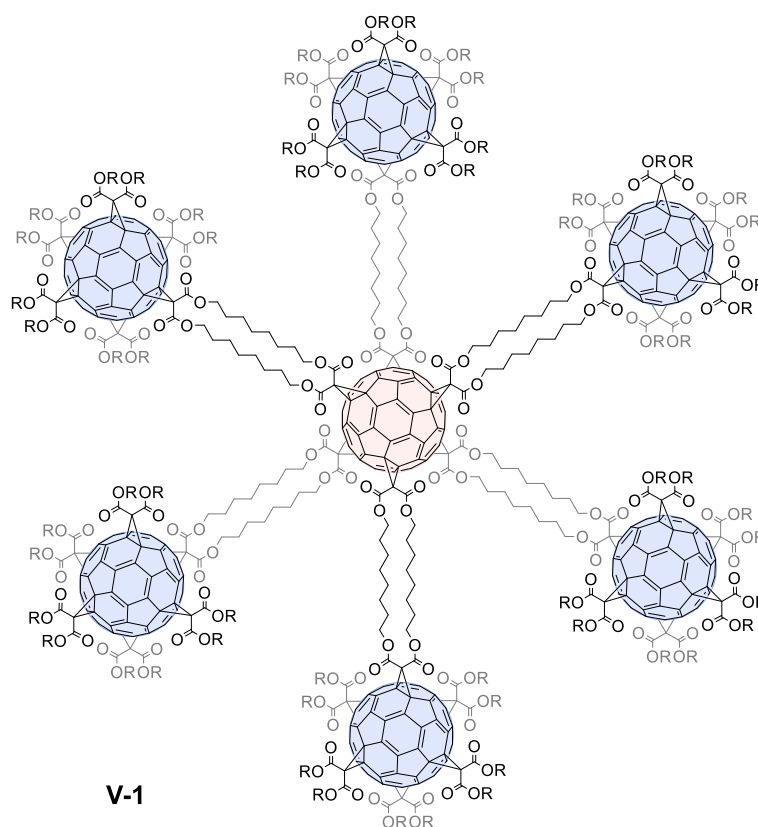


*Figure V-1* : Example of fullerodendrimer of C<sub>60</sub> hexa-adduct bearing two different type of addends.

The problem of accessibility of bulky hexa-adducts of C<sub>60</sub> has been resolved by Nierengarten's group.<sup>[8]</sup> Easily accessible C<sub>60</sub> hexa-adducts bearing 12 terminal alkyne or azide groups have been synthesized. Post-functionalization of these building blocks by copper-

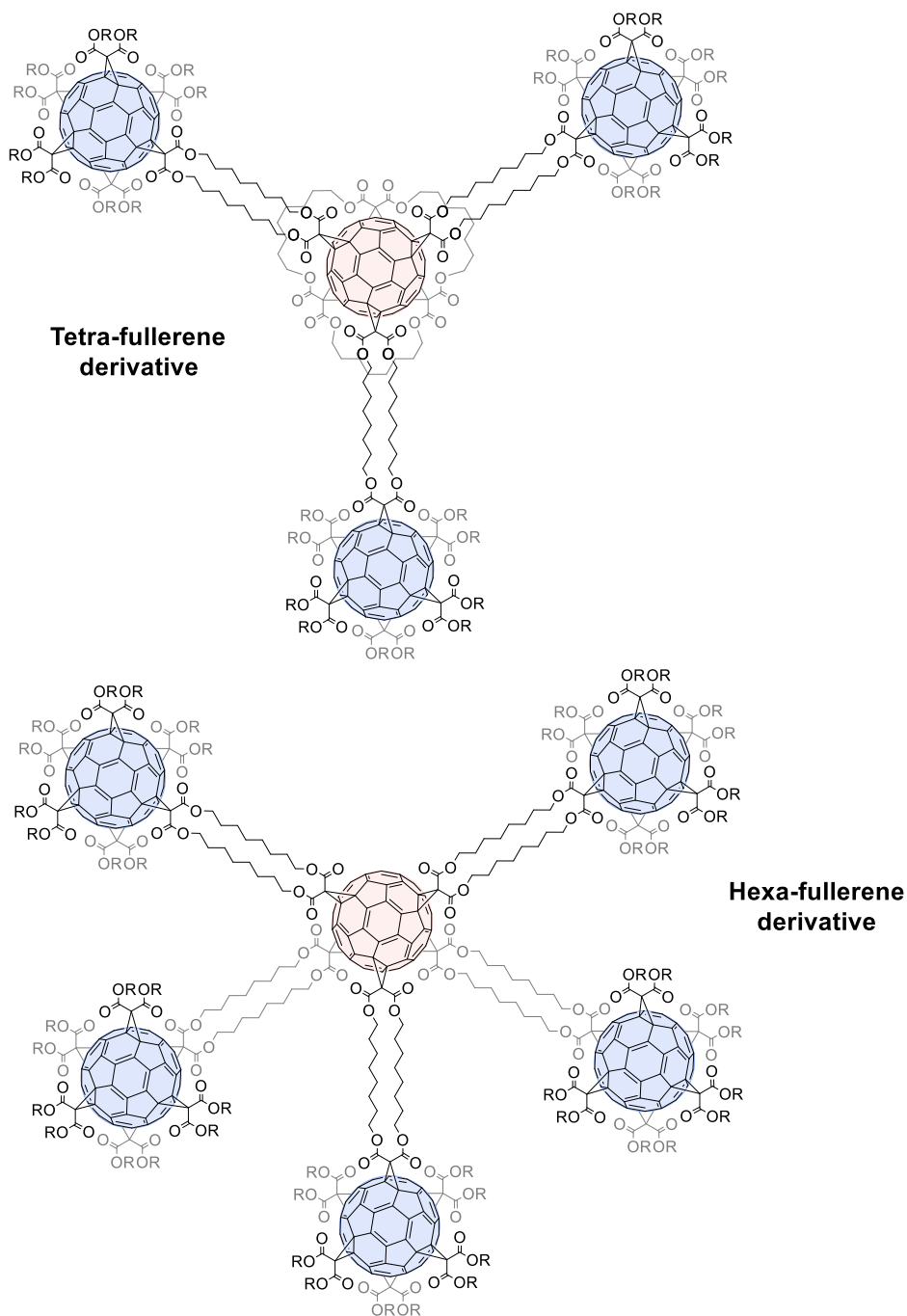
catalyzed alkyne-azide 1,3-dipolar cycloaddition (CuAAC) reactions has allowed the formation of C<sub>60</sub> hexa-adducts dendrimers in good yields.<sup>[9–13]</sup>

Among the fullerodendrimers, fullerene rich dendrimers have generated a particular interest.<sup>[6,7]</sup> The fullerene rich dendrimers reported in the literature are mainly focused on mono- or bis-functionalized C<sub>60</sub> at the periphery of the dendrimer limiting their applications to material science. The principal interest of such type of dendrimers is based on the physico-chemical properties of the C<sub>60</sub> sphere (the properties of pristine C<sub>60</sub> are kept in mono- and bis-functionalized C<sub>60</sub>). On the other hand, there are only few examples of fullerodendrimers with a C<sub>60</sub> core surrounded by multi-substituted C<sub>60</sub>. The synthesis of the hept fullerene **V-1** is the first example and was reported in 2009 by Hirsch and coworkers ([Figure V-2](#)).<sup>[14]</sup> The synthesis of this compound **V-1** was based on a [5:1] hexa-adduct of C<sub>60</sub> and on macrocyclic linkers. However, each step of the synthesis suffers of low yield and fastidious purification by the means of preparative HPLC is required.



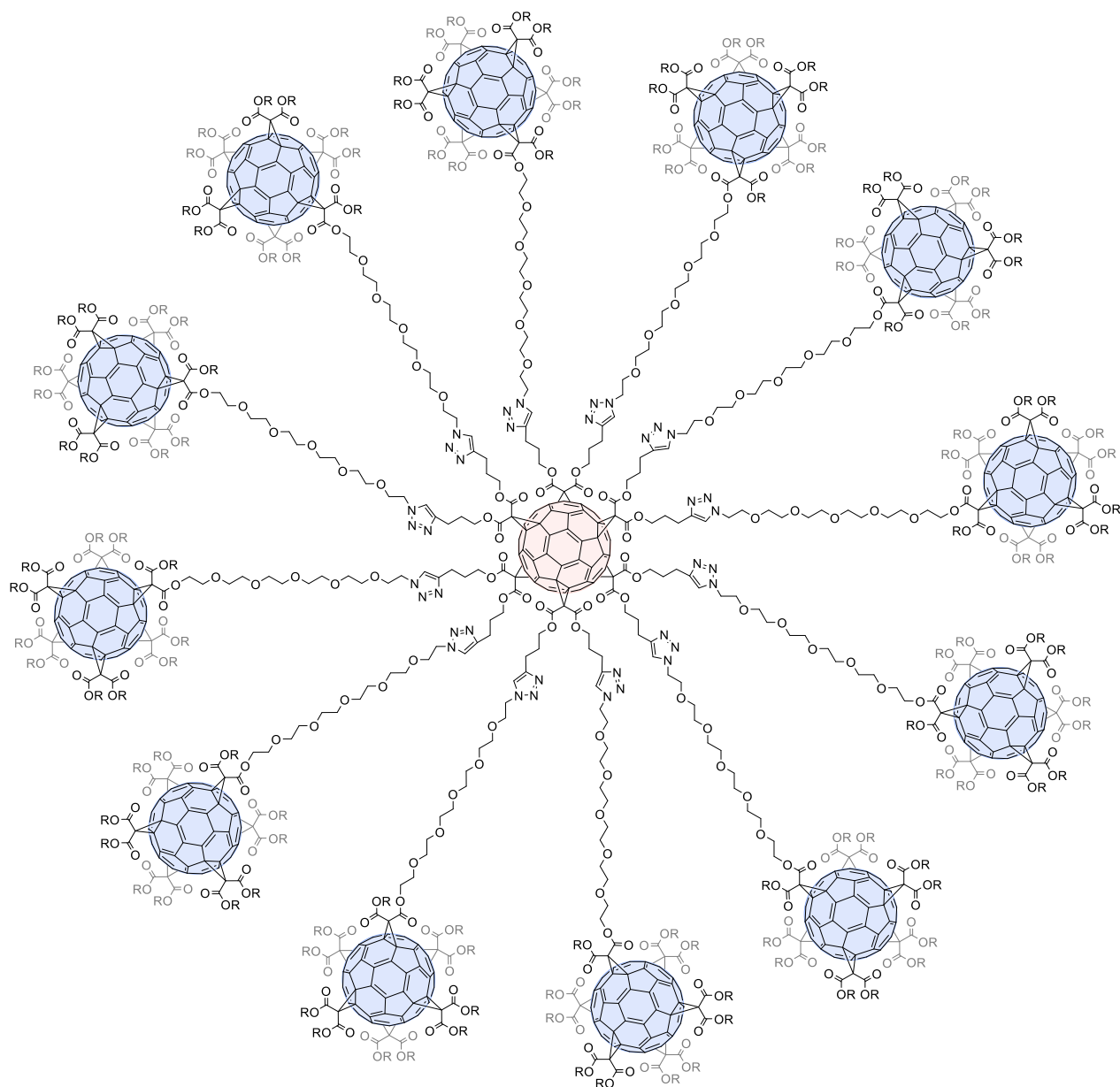
[Figure V-2](#) : First example of fullerodendrimer with a C<sub>60</sub> core surrounded by multi-substituted C<sub>60</sub>.

Later, Hirsch and coworker reported also the syntheses of tetra- and hexafullerenes with a precisely defined structure ([Figure V-3](#)).<sup>[15]</sup> In this case, a different approach was used for the synthesis of the key intermediate [5:1] hexa-adducts of C<sub>60</sub> which was isolated in much better yields. Furthermore, two other central building blocks were used. Nonetheless, the syntheses of the tetra- and hexafullerenes were subjected to the same synthetic difficulty as for the preparation of **V-1**.



**Figure V-3:** Structures of the tetrafullerene and hexafullerene derivatives.

Recently, Nierengarten's group has reported the syntheses of dendritic tridecafullerene derivatives (**Figure V-4**).<sup>[16,17]</sup> The tridecafullerene derivatives were obtained by the grafting of [5:1] hexa-adducts of  $C_{60}$  onto a central  $C_{60}$  hexa-adduct bearing 12 alkyne functions under copper catalyzed azide-alkyne cycloaddition conditions. These fullerodendrimers of first generation are surrounded by 120 or 132 peripheral functions. This synthetic strategy provides an efficient access to giant functional macromolecules in a minimum of synthetic steps.



*Figure V-4 : Dendritic tridecafullerene derivative synthesized by "click" reaction.*

The synthesis of fullerodendrimers with different types of functions disposed in a precisely defined structure at the periphery is still a challenging task. The regio- and stereoselective functionalization of C<sub>60</sub> and the subsequent post-functionalization of the fullerene derivatives are the main difficulties in order to have access to such type of fullerodendrimers.

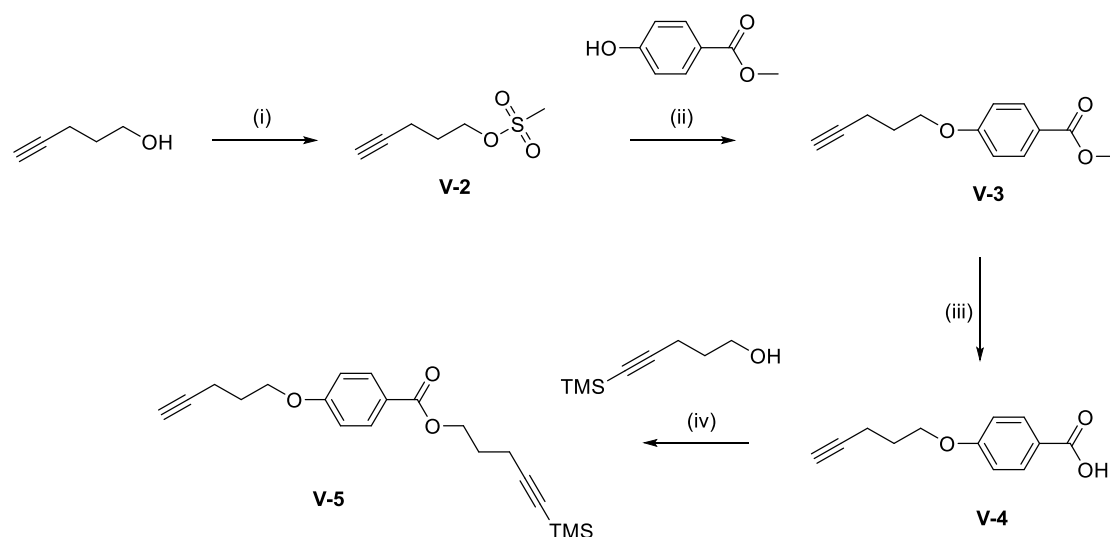
Based on the mixed C<sub>60</sub> hexa-adducts building blocks developed in the **Chapter IV**, the synthesis of 4 structurally new fullerodendrimers with two different types of functions at the periphery are presented in this chapter.

## 2. Preparation of tri-, tetra- and pentafullerene derivatives.

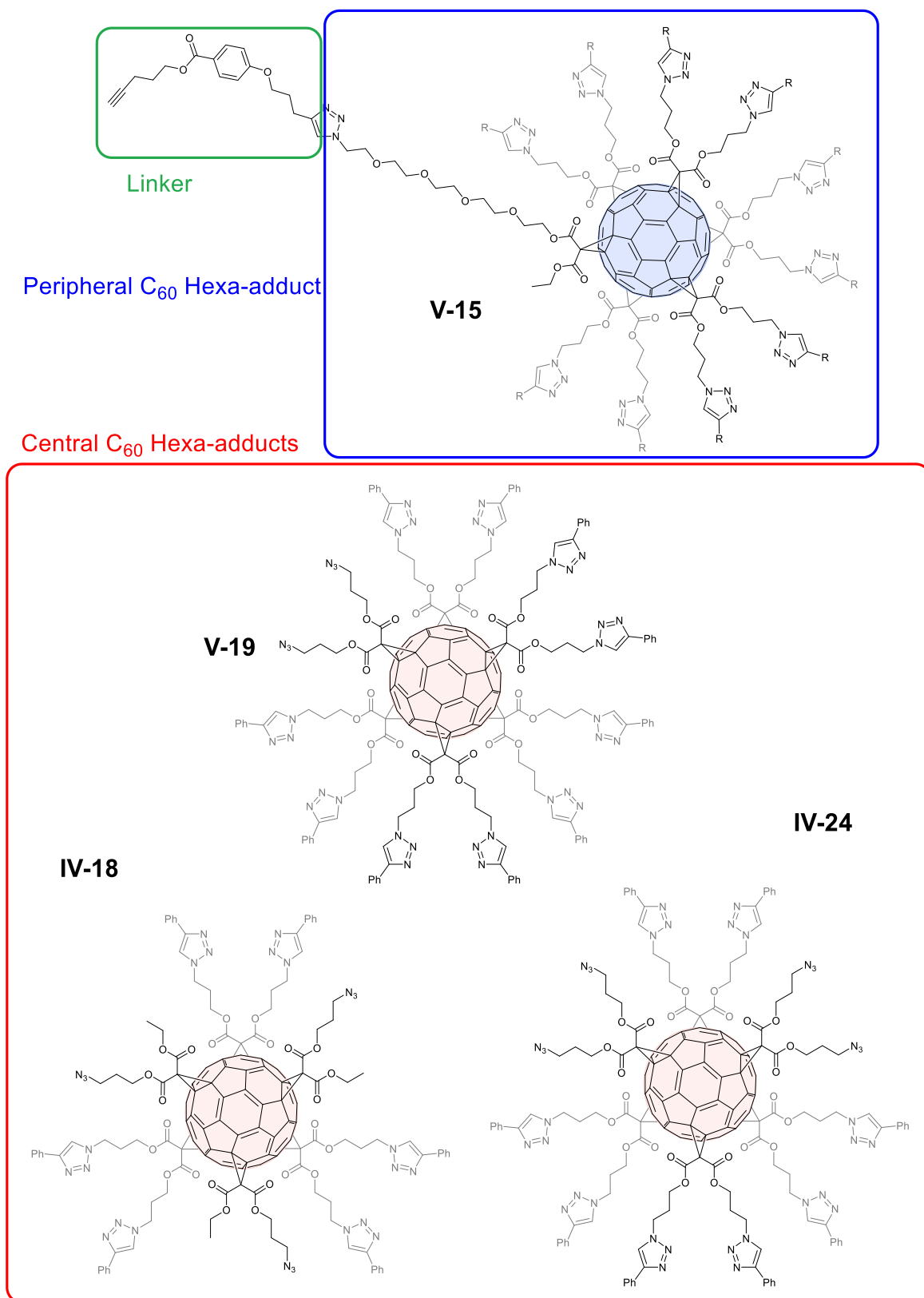
The synthesis of the tri-, tetra and pentafullerene derivatives **V-20-22** relies on the grafting of peripheral C<sub>60</sub> [5:1] hexa-adducts **V-15** onto the central mixed C<sub>60</sub> hexa-adducts building blocks **V-19**, **IV-18** and **IV-24** (Figure V-5). For synthetic reasons, the peripheral and the central C<sub>60</sub> hexa-adducts are both bearing azide functions. In order to allow the grafting of the peripheral hexa-adducts onto the central core, the synthesis of a dialkyne linker is therefore also required.

### 2.1 Preparation of the building blocks.

The synthesis of the dialkyne linker **V-5** is depicted in **Scheme V-1**. Treatment of 4-pentyn-1-ol with methanesulfonyl chloride (MeSO<sub>2</sub>Cl) in CH<sub>2</sub>Cl<sub>2</sub> in the presence of triethylamine gave compound **V-2**. Subsequent Williamson reaction of methyl 4-hydroxybenzoate with **V-2** afforded compound **V-3** in good yield (83%). Saponification of the methyl ester of **V-3** gave the corresponding carboxylic acid **V-4**. The key derivative **V-5** was then obtained by an esterification reaction of **V-4** with 5-(trimethylsilyl)-4-pentyn-1-ol under Steglich conditions.<sup>[18]</sup> A key feature of **V-5** is the protection of one alkyne function by a trimethylsilane group. The protected alkyne is inert under CuAAC reaction conditions and allows the functionalization of only one side of the linker **V-5**.



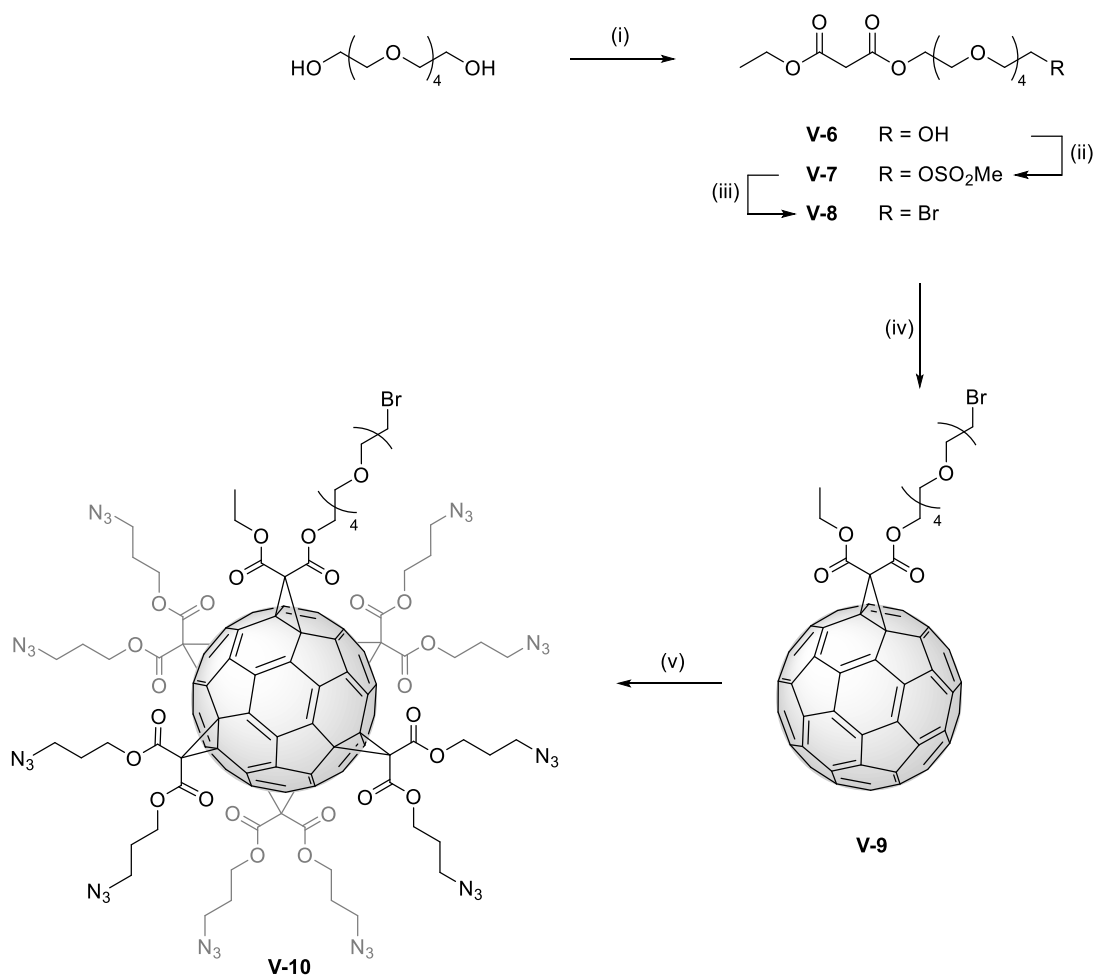
**Scheme V-1.** Reagents and conditions: (i) MeSO<sub>2</sub>Cl, trimethylamine, CH<sub>2</sub>Cl<sub>2</sub>, rt, 1 h (96%); (ii) K<sub>2</sub>CO<sub>3</sub>, crown 18C6 (cat.), acetone, reflux, 1.5 d (83%); (iii) KOH, THF/EtOH, reflux, 2.5 h (96%); (iv) DCC, DMAP, CH<sub>2</sub>Cl<sub>2</sub>, 0 °C to rt, 12 h (65%).



**Figure V-5:** C<sub>60</sub> hexa-adducts building blocks for the preparation the tri-, tetra and pentafullerene derivatives V-20-22.



The preparation of the peripheral C<sub>60</sub> [5:1] hexa-adduct **V-14** is depicted in **Scheme V-2**. Reaction of an excess of pentaethylene glycol with ethyl malonyl chloride in the presence of pyridine gave malonate **V-6** in 65% yield. Mesylation of **V-6** by treatment with MeSO<sub>2</sub>Cl/Et<sub>3</sub>N followed by a bromination reaction with LiBr afforded malonate **V-8**. Reaction of malonate **V-8** with C<sub>60</sub> in the presence of I<sub>2</sub> and DBU gave the mono-adduct **V-9** in 51% yield. Subsequent reaction of **V-9** with malonate **IV-12** (6.5 equiv.), CBr<sub>4</sub> (65 equiv.) and DBU (16 equiv.) in toluene gave hexa-adduct **V-10**. The “clicked” compounds **V-12a-b** were then prepared from **V-10** under typical CuAAC reaction conditions (**Scheme V-3**). Compounds **V-12a-b** have been characterized by NMR, UV-vis and IR spectroscopies and mass spectrometry. As previously reported for [5:1] C<sub>60</sub> hexa-adducts,<sup>[9,19,20]</sup> the <sup>13</sup>C NMR spectra of **V-12a-b** reveal the influence of the high local symmetry. Only one set of signals is observed for the five pseudoequivalent “clicked” malonates (**Figure V-6**). The signals of the unique malonate are also clearly distinguishable.



**Scheme V-2.** Reagents and conditions: (i) Ethyl malonyl chloride, pyridine, CH<sub>2</sub>Cl<sub>2</sub>, rt, 2 h (65%); (ii) MeSO<sub>2</sub>Cl, trimethylamine, CH<sub>2</sub>Cl<sub>2</sub>, rt, 1 h (91%); (iii) LiBr, THF, 60 °C, 4 h (87%); (iv) C<sub>60</sub>, I<sub>2</sub>, DBU, PhMe, rt, 1 h (51%); (v) **IV-12**, CBr<sub>4</sub>, DBU, PhMe, rt, 12 h (38%).

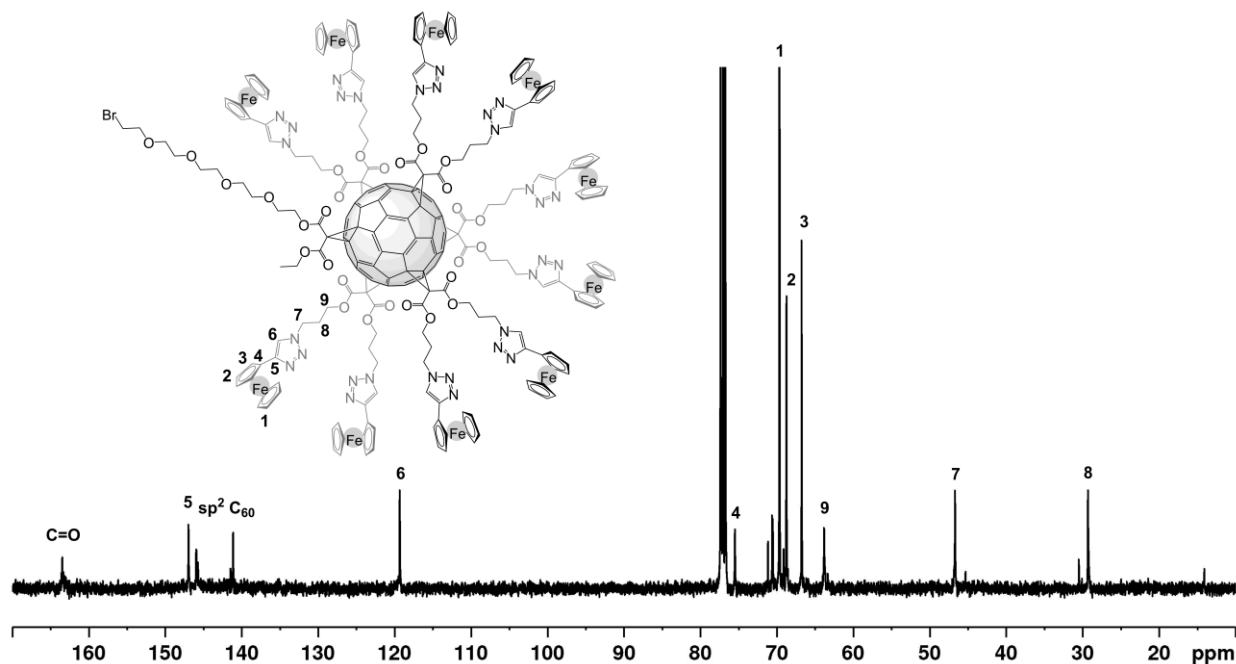
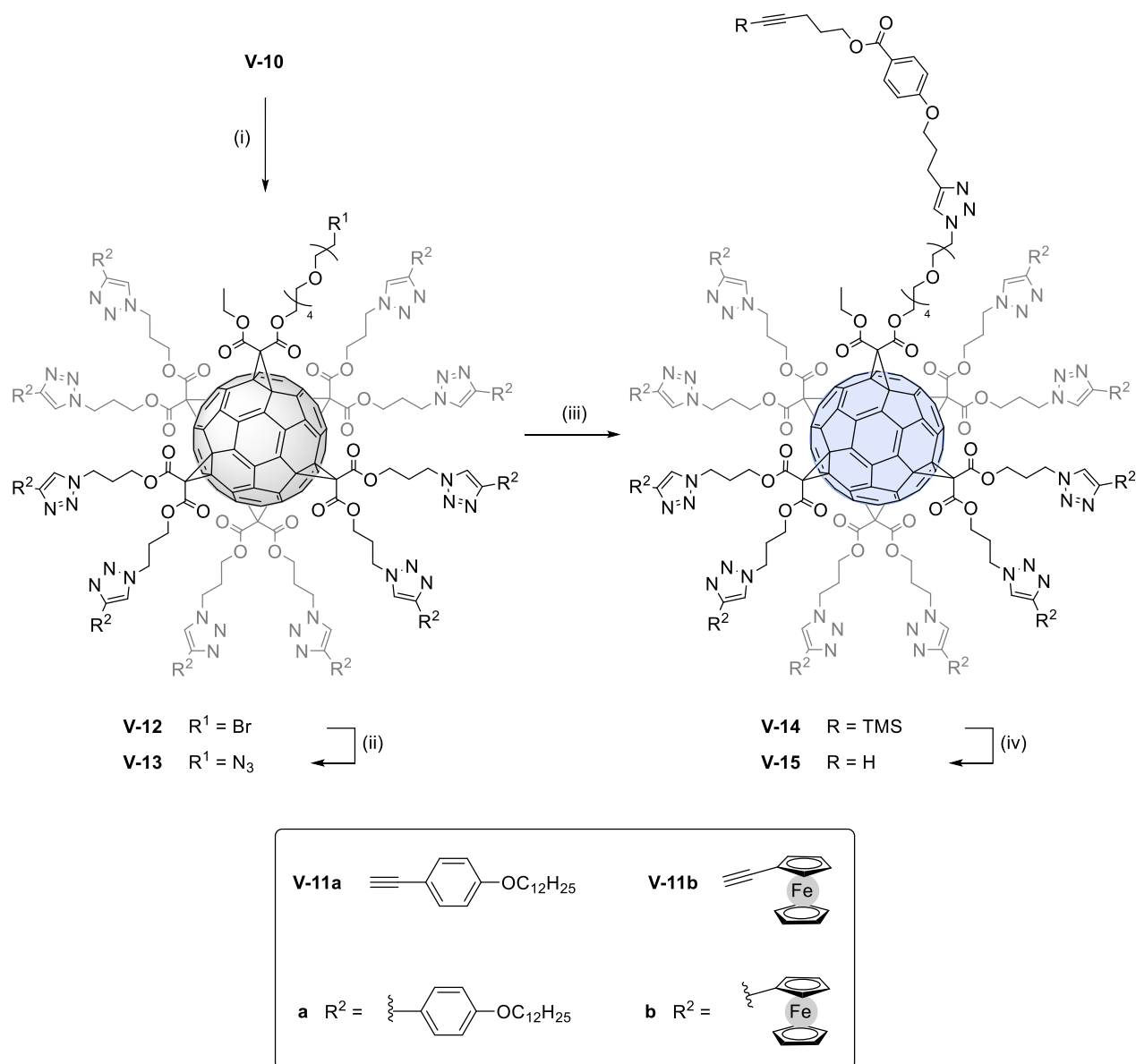


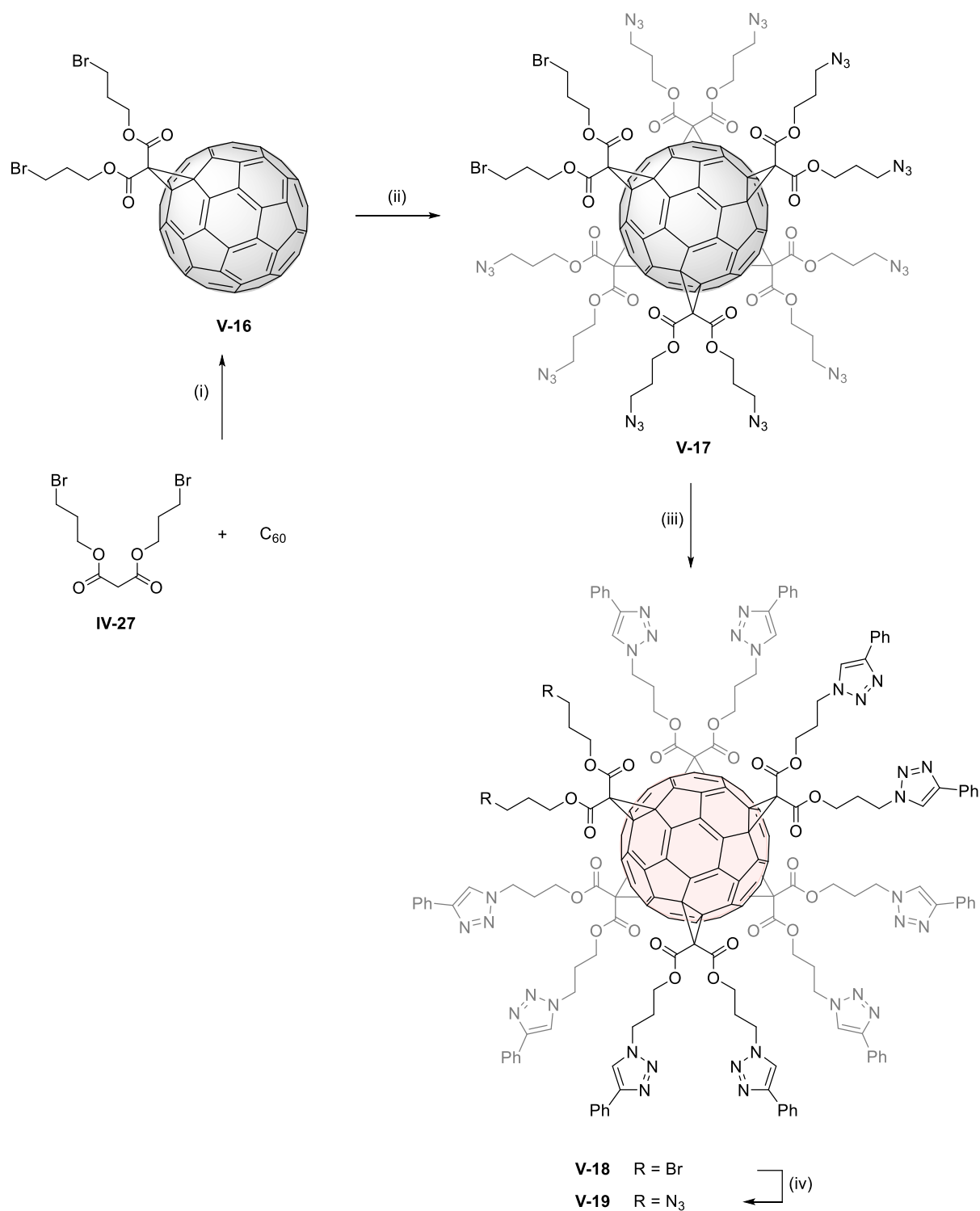
Figure V-6. <sup>13</sup>C NMR spectrum (100 MHz, CDCl<sub>3</sub>) of **V-12b** showing the influence of the high local symmetry.

The key derivatives **V-15a-b** were obtained in 3 steps from compound **V-12a-b** (Scheme V-3). Conversion of the bromides was carried out by treatment of **V-12a-b** with sodium azide in DMF to afford the corresponding azides (**V-13a-b**). Reaction of **V-13a-b** with the dialkyne linker **V-5** under CuAAC reaction conditions gave compounds **V-14a-b** in very good yields (99% and 88%, respectively). Finally, the deprotection of the trimethylsilyl group (TMS) was performed by treatment of **V-14a-b** with tetrabutylammonium fluoride (TBAF) in THF at 0 °C to afford the key derivatives **V-15a-b**. In both cases, the removal of the TMS group was confirmed by NMR and IR spectroscopies as well as by mass spectrometry.



**Scheme V-3.** Reagents and conditions: (i) **V-11a** or **b**, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (**V-12a**: 97%, **V-12b**: 87%); (ii) NaN<sub>3</sub>, DMF, rt, 12 h (**V-13a**: 93%, **V-13b**: 94%); (iii) **V-5**, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 2.5 h (**V-14a**: 99%, **V-14b**: 88%); (iv) TBAF, THF, 0 °C, 20 min (**V-15a**: 99%, **V-15b**: 92%).

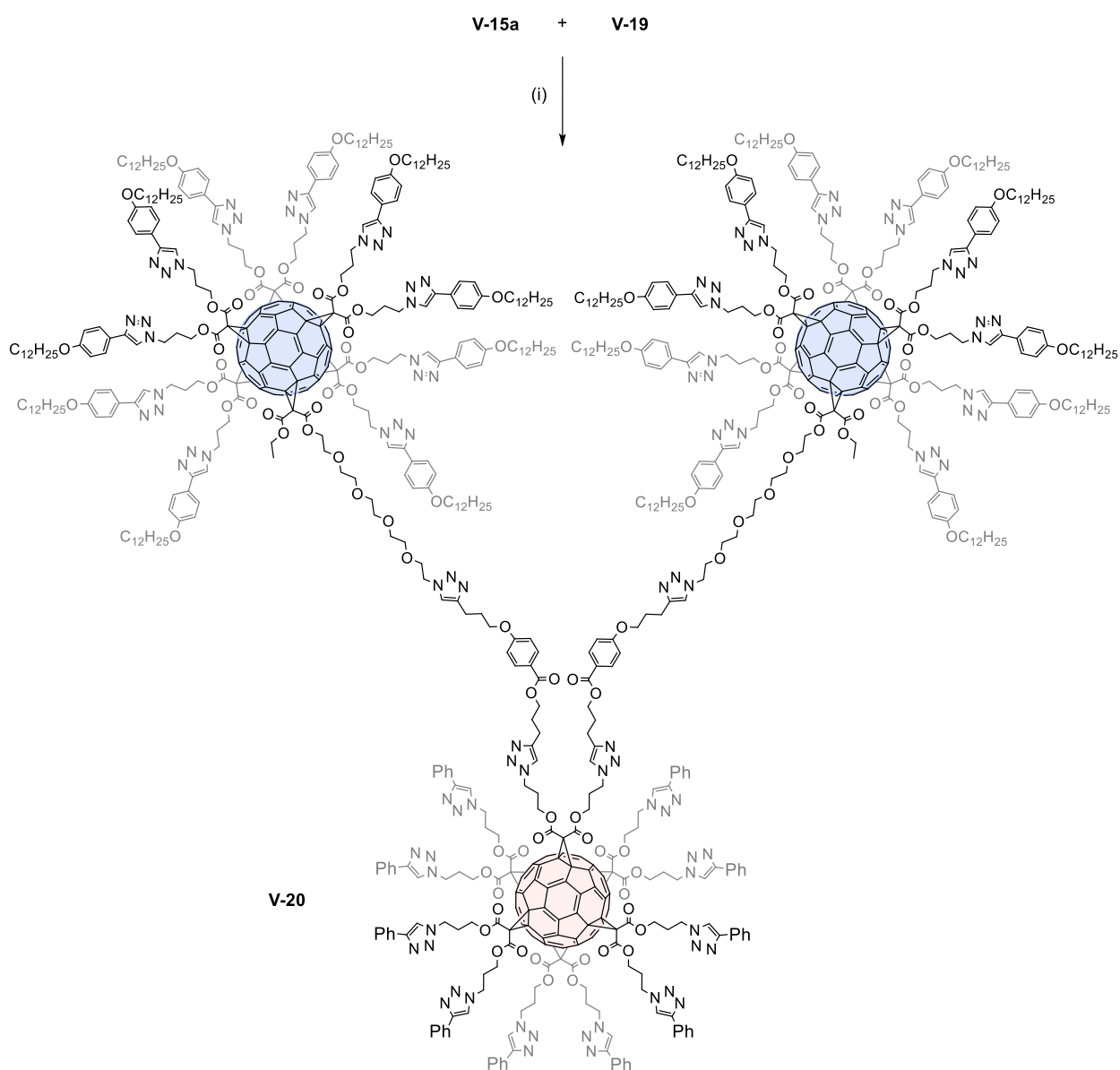
The synthesis of **V-19** is depicted in **Scheme V-4**. Reaction of the malonate **IV-27** with C<sub>60</sub> in the presence of I<sub>2</sub> and DBU afforded the mono-adduct **V-16** in 51% yield. Subsequent reaction of **V-16** with the malonate **IV-12** (7 equiv.), CBr<sub>4</sub> (70 equiv.) and DBU (17 equiv.) in toluene at room temperature gave the hexa-adduct **V-17**. Reaction of **V-17** with phenylacetylene under typical conditions of CuAAC reactions gave the “click” compound **V-18** in a good yield (79%). Compound **V-18** has been characterized by NMR, UV-vis and IR spectroscopies. In addition, the MALDI-TOF mass spectrum was also consistent with the proposed structure. Finally, the bromide functions in **V-18** were converted into azides to afford the mixed C<sub>60</sub> hexa-adduct building blocks **V-19**.



**Scheme V-4.** Reagents and conditions: (i) I<sub>2</sub>, DBU, PhMe, rt, 1 h (51%); (ii) **IV-12**, CBr<sub>4</sub>, DBU, PhMe, 12 h (55%); (iii) Phenylacetylene, CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (79%); (iv) NaN<sub>3</sub>, DMF, rt, 12 h (93%).

## 2.2 Synthesis of tris-, tetra- and pentafullerene derivatives.

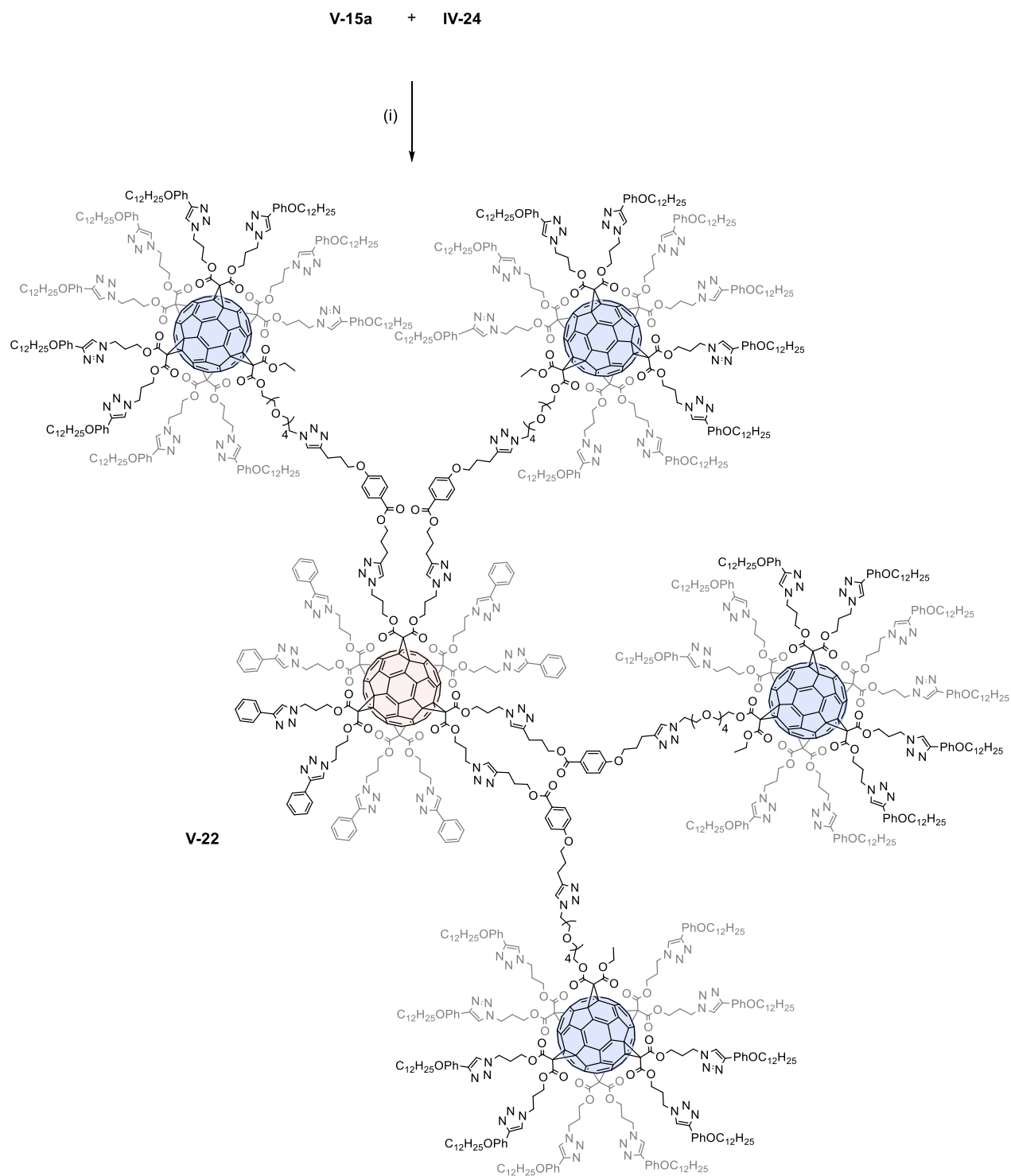
Based on the previously prepared building blocks, the syntheses of the trisfullerene derivative **V-20** (Scheme V-5), the tetrafullerene derivative **V-21** (Scheme V-6) and the pentafullerene derivative **V-22** (Scheme V-7) were achieved. Reaction of the peripheral C<sub>60</sub> [5:1] hexa-adducts **V-15a** with the appropriated mixed C<sub>60</sub> hexa-adduct building block (**V-19**, **IV-18** or **IV-24**) under CuAAC reaction conditions afforded the compounds **V-20-22**. Purification of **V-20-22** were achieved by column chromatography on silica gel followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>).



**Scheme V-5.** Reagents and conditions: (i) CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (56%).



**Scheme V-6.** Reagents and conditions: (i) CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (62%).



**Scheme V-7.** Reagents and conditions: (i) CuSO<sub>4</sub>·5H<sub>2</sub>O, sodium ascorbate, CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O, 30 °C, 12 h (68%).

Compounds **V-20-22** have been characterized by NMR, UV-vis and IR spectroscopies and mass spectrometry. The <sup>1</sup>H NMR spectrum of **V-22** is showed in [Figure V-7](#) as typical example. The appearance of a second resonance between  $\delta = 2.8-2.9$  ppm relative to the CH<sub>2</sub>C<sub>triazole</sub> signal indicates that compound **V-22** is “clicked” on compound **IV-24**. The integration ratio of the signals of each different protons is also compatible with the proposed structure. Furthermore, the <sup>1</sup>H NMR spectrum reveals the absence of resonance at  $\delta = 1.9$  and 3.3 ppm relative to the CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub> parts of **IV-24** and at  $\delta = 2.0$  ppm relative to the CH<sub>2</sub>CH<sub>2</sub>C≡CH parts of **V-15a**. These latter observations are also confirmed by the IR data with the absence of signal for the azide (2098 cm<sup>-1</sup>) and for the alkyne (3315 cm<sup>-1</sup>).

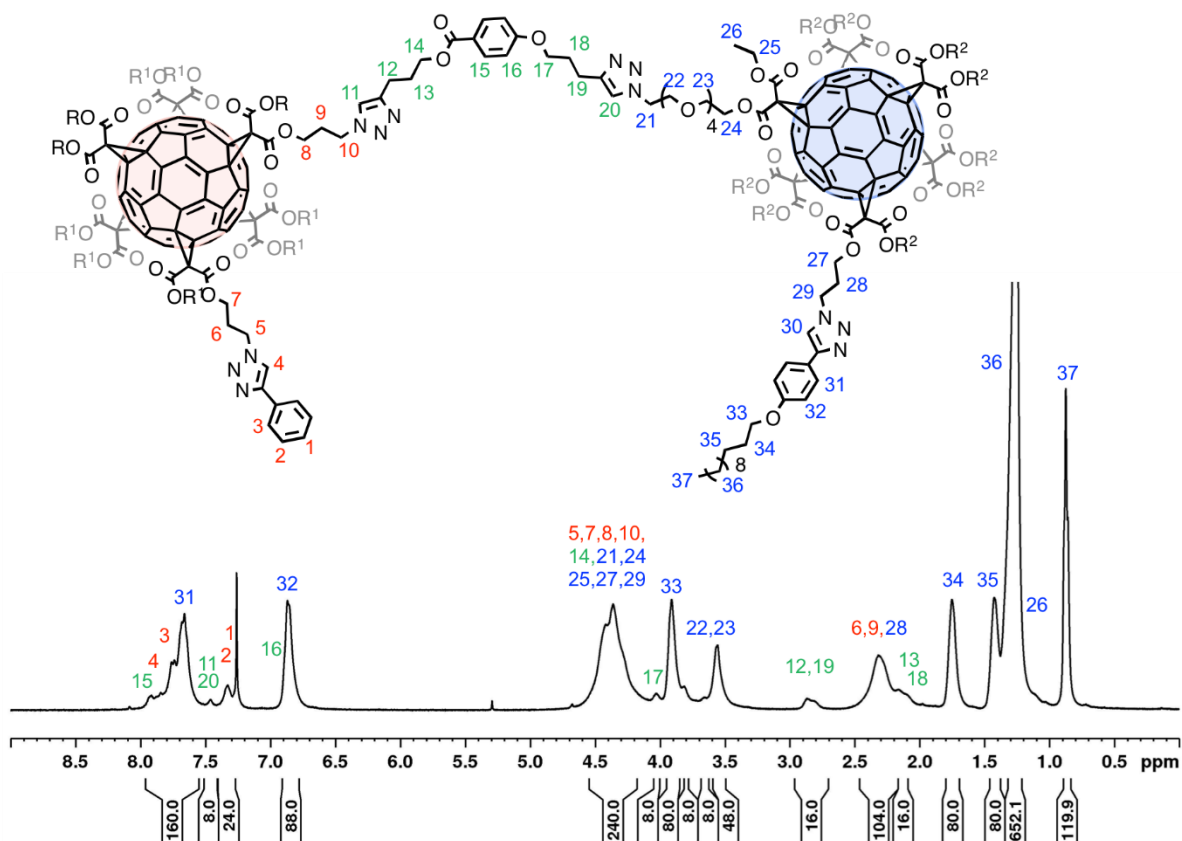


Figure V-7.: <sup>1</sup>H NMR spectrum (400MHz, CDCl<sub>3</sub>) of **V-22**.

In addition, the MALDI-TOF mass spectra were consistent with the proposed structures. Due to the high molecular weight, the signals are broad and thus the resolution of the spectra are not optimum. Furthermore, high level of fragmentation is also observed. In the case of compounds **V-20** and **V-22**, the molecular ion peak could not be measured precisely (**V-20**:  $m/z = 14489$  ([M]<sup>+</sup>, calcd for C<sub>868</sub>H<sub>938</sub>O<sub>106</sub>N<sub>102</sub>: 14495); **V-22**:  $m/z \approx 25440$  ([M]<sup>+</sup>, calcd for C<sub>1526</sub>H<sub>1732</sub>O<sub>188</sub>N<sub>168</sub>: 25434)). For compound **V-21**, the molecular ion peak is not detected. Nonetheless, characteristic fragmentation resulting from ester hydrolysis followed by decarboxylation and retro-Bingel reactions on the peripheral C<sub>60</sub> hexa-adducts are observed on the three mass spectra ([Figure V-8](#)).<sup>[21,22]</sup> All these observations are proofs of the formation of compounds **V-20-22**.



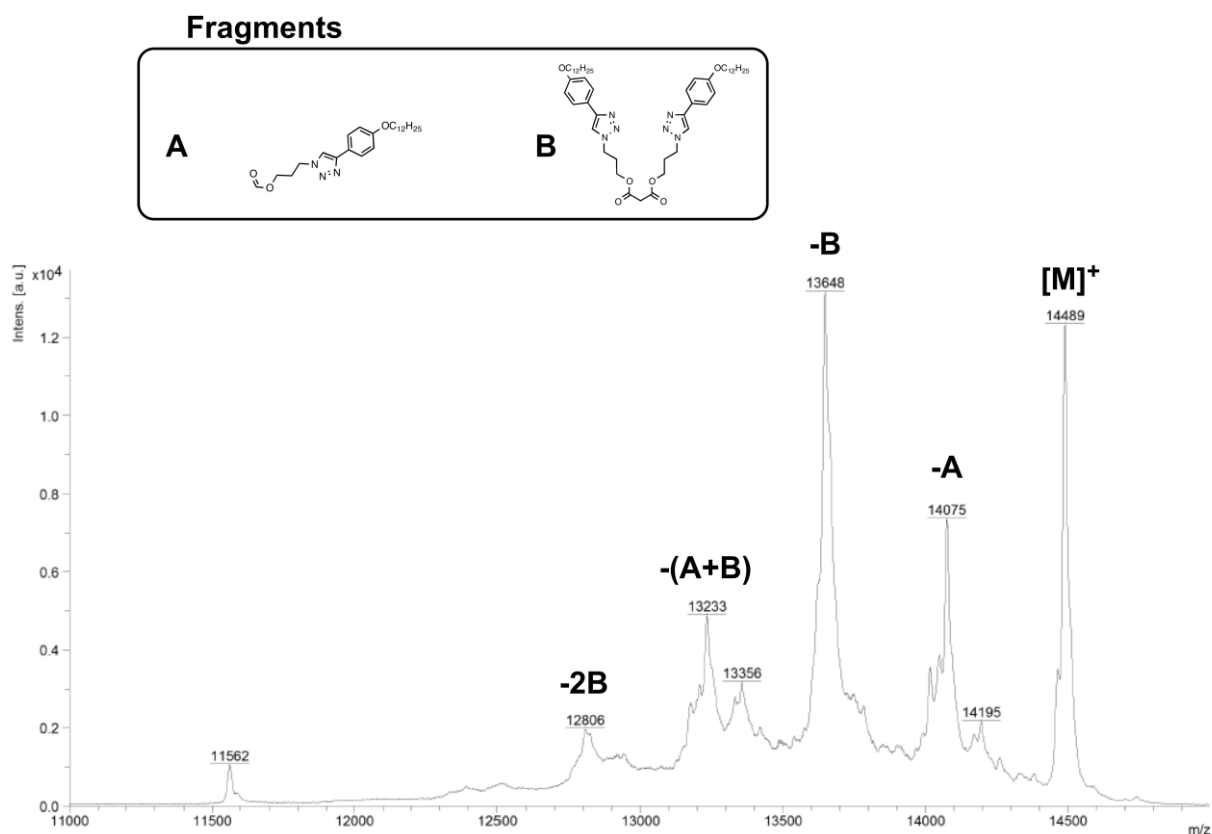


Figure V-8 : MALDI-TOF mass spectrum of **V-20** showing the characteristic fragmentations.

Compounds **V-20-22** are decorated with two types of functions in a 20:10, 30:6 and 40:8 ratio, respectively. Each peripheral C<sub>60</sub> hexa-adduct bears 10 dodecoxyphenyl groups and the central C<sub>60</sub> hexa-adducts 6 to 10 phenyl groups. Compounds **V-20-22** represent new examples of multi- and poly- functional molecules synthesized with a perfect control of the regiochemistry.

### 3. Giant mixed C<sub>60</sub> hexa-adducts dendrimer.

The synthesis of the giant mixed tridecafullerene **V-25** relies on the sequential grafting of two different peripheral C<sub>60</sub> [5:1] hexa-adducts (**V-15a** and **V-15b**) onto a central mixed C<sub>60</sub> hexa-adduct building block (**IV-16**). Reaction of **V-15a** with **IV-16** under CuAAC reaction conditions afforded the nonafullerene derivative **V-23** (**Scheme V-8**). The <sup>1</sup>H NMR spectrum of **V-23** reveals the same characteristic features as described previously for compound **V-22**. In addition, the IR data show the absence of unreacted azide (2098 cm<sup>-1</sup>) and alkyne (3315 cm<sup>-1</sup>) residues. Treatment of **V-23** with sodium azide in DMF was next performed to afford compound **V-24**. The total conversion into azide functions was confirmed by the change in chemical shift for the terminal methylene group: from  $\delta = 3.41$  ppm for the CH<sub>2</sub>Br signals to  $\delta = 3.35$  ppm for the CH<sub>2</sub>N<sub>3</sub> signals. Finally, **V-25** was obtained by a second CuAAC reaction of **V-24** with **V-15b** (**Scheme V-9**). The <sup>1</sup>H NMR spectrum of **V-25** is showed in [Figure V-9](#). The signals relative to the substituents of the two different peripheral C<sub>60</sub> hexa-adducts are clearly distinguishable. Furthermore, the resonances attributed to the aromatic protons of the 1,2,3-triazole and the CH<sub>2</sub>C<sub>triazole</sub> of the linker sub-units are observed at  $\delta = 7.45$  and 2.85 ppm, respectively. The integration ratio of each different signal is also in agreement with the proposed structure. The <sup>13</sup>C NMR spectrum of **V-25** shows the strong influence of the high local symmetry ([Figure V-9](#)). The differentiation is only observed for the terminal sub-units grafted onto the peripheral C<sub>60</sub> hexa-adducts. The characteristic signals of [C<sub>18,27</sub>] and [C<sub>19,26</sub>] of the linker subunits are detected at  $\delta = 122.0$  and 22.2 ppm. The signal of [C<sub>17</sub>] is also detected at  $\delta = 50.1$  ppm. The MALDI-TOF experiment was not suited for compound **V-25** and only broad signals were observed. Nonetheless, these signals are in the expected molecular weight range.

Compound **V-25** is a first generation dendrimer possessing 120 peripheral functions. 80 dodecoxyphenyl groups are brought by 8 peripheral C<sub>60</sub> hexa-adducts with the first CuAAC reaction and 40 ferrocene groups by 4 other peripheral C<sub>60</sub> hexa-adducts with the second CuAAC reaction. Compound **V-25** is the first example of giant multi-functionalized fullerodendrimer synthesized with the control of the regiochemistry.





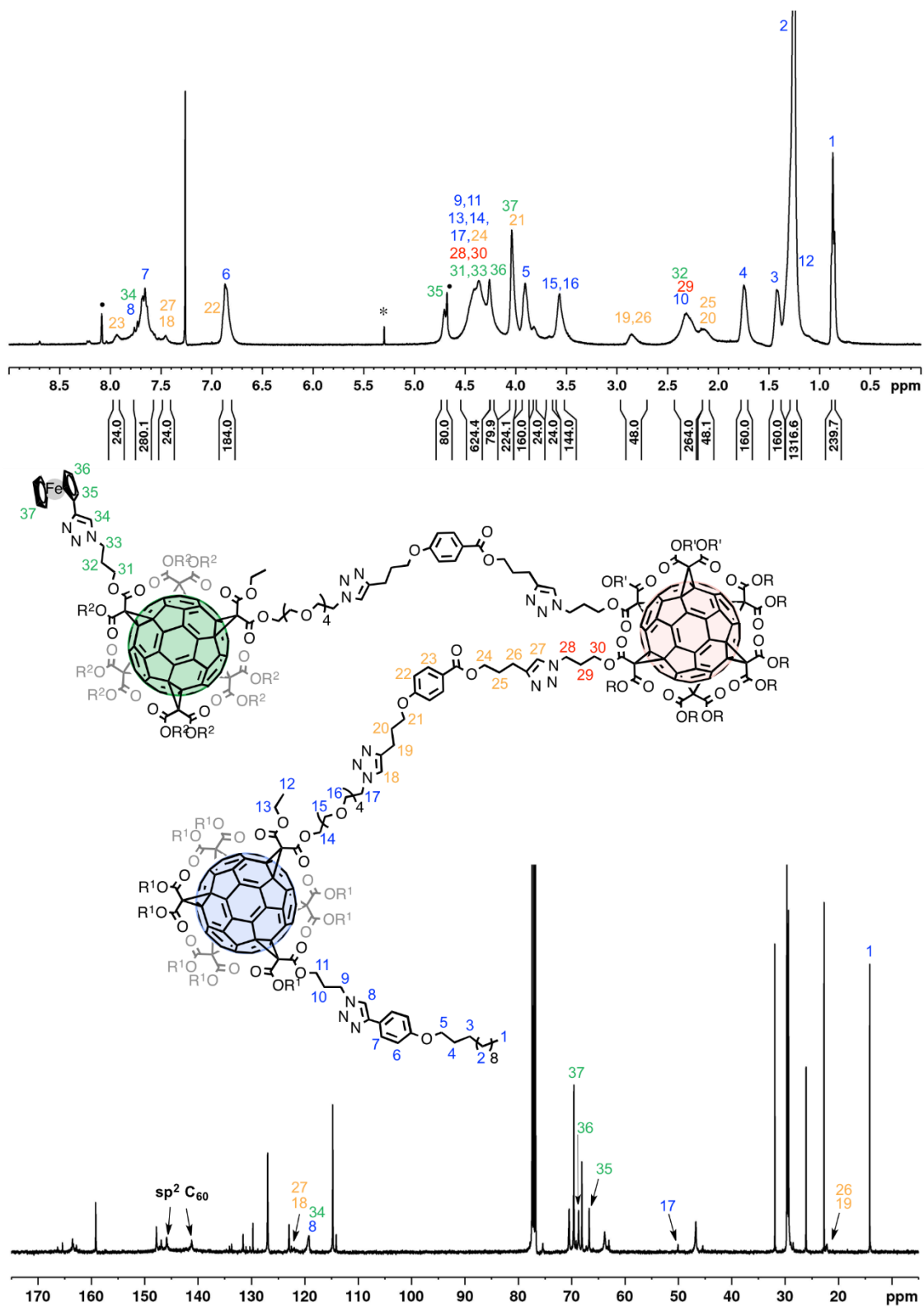


Figure V-9 : (top) <sup>1</sup>H NMR spectrum (400MHz, CDCl<sub>3</sub>) of V-25, \* CH<sub>2</sub>Cl<sub>2</sub>, • impurities; (bottom) <sup>13</sup>C NMR spectrum (100MHz, CDCl<sub>3</sub>) of V-25.

## 4. Conclusion.

The synthesis of fullerodendrimers based on C<sub>60</sub> mixed hexa-adduct building blocks has been achieved. The grafting of the peripheral C<sub>60</sub> [5:1] hexa-adducts onto a central C<sub>60</sub> hexa-adduct has been efficiently performed under CuAAC reaction conditions. Two defined function zones are thus generated. Different ratios of the functional groups have been thus obtained (20:10, 30:6, 40:8 and 80:40). Based on our different C<sub>60</sub> hexa-adducts building blocks, the controlled elaboration of new nanomaterials and bioactive molecules should become easily accessible.

## 5. Bibliography.

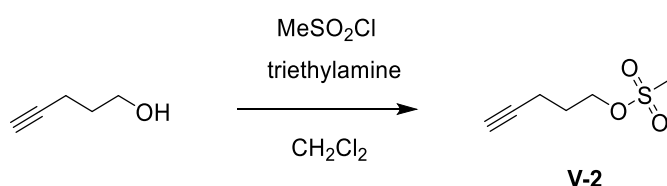
- [1] G. R. Newkome, C. N. Moorefield, F. Vögtle, in *Dendrimers Dendrons Concepts Synth. Appl.*, Wiley-VCH Verlag GmbH & Co. KGaA, **2001**.
- [2] J. M. J. Fréchet, D. A. Tomalia, Eds. , in *Dendrimers Dendritic Polym.*, John Wiley & Sons, Ltd, **2001**.
- [3] K. L. Wooley, C. J. Hawker, J. M. J. Frechet, F. Wudl, G. Srdanov, S. Shi, C. Li, M. Kao, *J. Am. Chem. Soc.* **1993**, *115*, 9836–9837.
- [4] A. Hirsch, O. Vostrowsky, in *Dendrimers IV* (Eds.: P.D.F. Vögtle, D.C.A. Schalley), Springer Berlin Heidelberg, **2001**, pp. 51–93.
- [5] A. Hirsch, O. Vostrowsky, *Eur. J. Org. Chem.* **2001**, *2001*, 829–848.
- [6] M. Holler, J. Nierengarten, *Aust. J. Chem.* **2009**, *62*, 605–623.
- [7] U. Hahn, F. Vögtle, J.-F. Nierengarten, *Polymers* **2012**, *4*, 501–538.
- [8] J. Iehl, R. P. de Freitas, B. Delavaux-Nicot, J.-F. Nierengarten, *Chem. Commun.* **2008**, 2450–2452.
- [9] J. Iehl, J.-F. Nierengarten, *Chem. – Eur. J.* **2009**, *15*, 7306–7309.
- [10] M. Sánchez-Navarro, A. Muñoz, B. M. Illescas, J. Rojo, N. Martín, *Chem. – Eur. J.* **2011**, *17*, 766–769.
- [11] D. Sigwalt, M. Holler, J. Iehl, J.-F. Nierengarten, M. Nothisen, E. Morin, J.-S. Remy, *Chem. Commun.* **2011**, *47*, 4640–4642.
- [12] K. Buffet, E. Gillon, M. Holler, J.-F. Nierengarten, A. Imberty, S. P. Vincent, *Org. Biomol. Chem.* **2015**, *13*, 6482–6492.
- [13] S. Guerra, J. Iehl, M. Holler, M. Peterca, D. A. Wilson, B. E. Partridge, S. Zhang, R. Deschenaux, J.-F. Nierengarten, V. Percec, *Chem. Sci.* **2015**, *6*, 3393–3401.
- [14] P. Witte, F. Hörmann, A. Hirsch, *Chem. – Eur. J.* **2009**, *15*, 7423–7433.
- [15] F. Hörmann, A. Hirsch, *Chem. – Eur. J.* **2013**, *19*, 3188–3197.
- [16] D. Sigwalt, R. Caballero, M. Holler, J.-M. Strub, A. Van Dorsselaer, J.-F. Nierengarten, *Eur. J. Org. Chem.* **2016**, *2016*, 2882–2887.

- [17] A. Muñoz, D. Sigwalt, B. M. Illescas, J. Luczkowiak, L. Rodríguez-Pérez, I. Nierengarten, M. Holler, J.-S. Remy, K. Buffet, S. P. Vincent, et al., *Nat. Chem.* **2016**, *8*, 50–57.
- [18] B. Neises, W. Steglich, *Angew. Chem.* **1978**, *90*, 556–557.
- [19] A. Herzog, A. Hirsch, O. Vostrowsky, *Eur. J. Org. Chem.* **2000**, *2000*, 171–180.
- [20] P. Fortgang, E. Maisonhaute, C. Amatore, B. Delavaux-Nicot, J. Iehl, J.-F. Nierengarten, *Angew. Chem. Int. Ed.* **2011**, *50*, 2364–2367.
- [21] M. Durka, K. Buffet, J. Iehl, M. Holler, J.-F. Nierengarten, S. P. Vincent, *Chem. – Eur. J.* **2012**, *18*, 641–651.
- [22] J. Li, L. C. Nye, L. K. Wasserthal, C. Vinh, R. W. Kirschbaum, I. Ivanović-Burmazović, A. Hirsch, T. Drewello, *Eur. J. Org. Chem.* **2015**, *2015*, 2282–2290.

## 6. Experimental part.

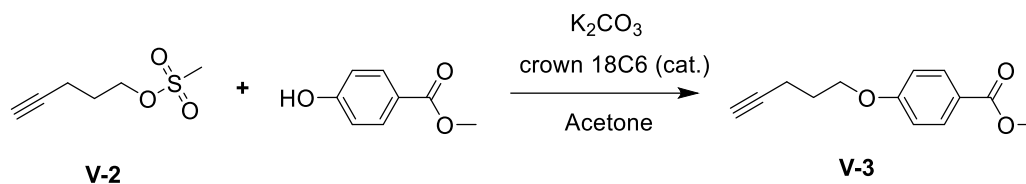
**General:** Reagents and solvents were purchased as reagent grade and used without further purification. All reactions were performed in standard glassware under an inert Ar atmosphere. Evaporation and concentration were done at water aspirator pressure and drying in vacuo at 10<sup>-2</sup> Torr. Column chromatography: silica gel 60 (230-400 mesh, 0.040-0.063 mm) was purchased from E. Merck. Thin Layer Chromatography (TLC) was performed on aluminum sheets coated with silica gel 60 F<sub>254</sub> purchased from E. Merck. IR spectra (cm<sup>-1</sup>) were recorded on a Perkin–Elmer Spectrum One Spectrophotometer. UV/Visible spectra were recorded on a Perkin-Elmer Lambda 35 UV/Vis Spectrometer. NMR spectra were recorded on a Bruker AC 300 or AC 400 with solvent peaks as reference at 25°C. MALDI-TOF-MS were obtained on a Bruker ULTRAFLEX TOF/TOF mass spectrometer with a dithranol matrix.

### Synthesis.

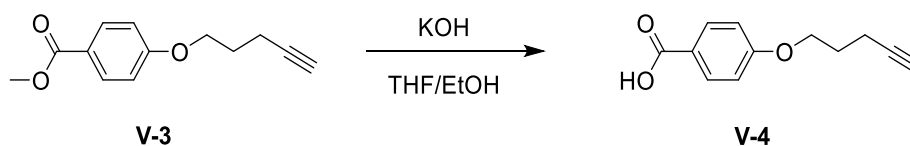


**Compound V-2:** Methanesulfonyl chloride (5 mL, 64 mmol) was added to a solution of 4-pentyn-1-ol (3 mL, 32 mmol) and triethylamine (9 mL, 64 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (100 mL). The resulting mixture was stirred at room temperature for 1 h, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5 to CH<sub>2</sub>Cl<sub>2</sub>) yielded **V-2** (5 g, 30.8 mmol, 96%) as a white solid. IR (neat):  $\nu = 3287$  (C≡CH), 2119 (C≡C) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.36$  (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>O), 3.03 (s, 3 H, OSO<sub>2</sub>CH<sub>3</sub>), 2.37 (dt, <sup>3</sup>J = 6

Hz &  $^4J = 2$  Hz, 2 H, C $\equiv$ CCH<sub>2</sub>CH<sub>2</sub>), 2.01 (t,  $^4J = 2$  Hz, 1 H, C $\equiv$ CH), 1.97 (quint.,  $^3J = 6$  Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 82.1, 69.8, 68.2, 37.3, 27.8, 14.7$  ppm.

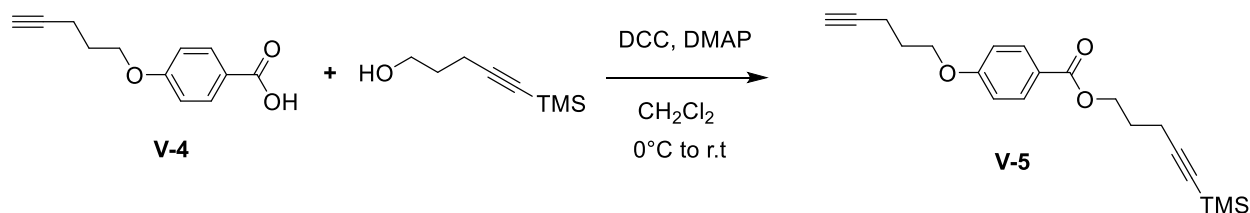


**Compound V-3:** **V-2** (5.00 g, 30.8 mmol) was added to a solution of methyl 4-hydroxybenzoate (4.69 g, 30.8 mmol), K<sub>2</sub>CO<sub>3</sub> (17 g, 123 mmol) and crown 18C6 (cat.) in acetone (250 mL). The resulting mixture was stirred under reflux for 1.5 days, then the salts were filtered and the product was extracted with Et<sub>2</sub>O, washed with H<sub>2</sub>O and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 7:3) yielded **V-3** (5.57 g, 25.5 mmol, 83%) as a white solid. IR (neat):  $\nu = 3296$  (C $\equiv$ CH), 2119 (C $\equiv$ C), 1712 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.98$  (d,  $^3J = 8$  Hz, 2 H, ArH<sub>o</sub>), 6.91 (d,  $^3J = 8$  Hz, 2 H, ArH<sub>m</sub>), 4.12 (t,  $^3J = 6$  Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>OAr), 3.88 (s, 3 H, COOCH<sub>3</sub>), 2.41 (dt,  $^3J = 6$  Hz &  $^4J = 2$  Hz, 2 H, C $\equiv$ CCH<sub>2</sub>CH<sub>2</sub>), 2.02 (quint.,  $^3J = 6$  Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.98 (t,  $^4J = 2$  Hz, 1 H, C $\equiv$ CH) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.8, 162.7, 131.6, 122.6, 114.6, 83.2, 69.1, 66.3, 51.8, 28.0, 15.1$  ppm.

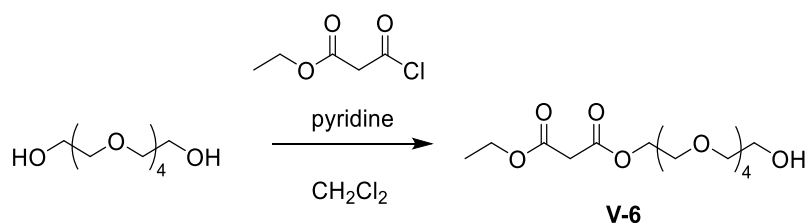


**Compound V-4:** A solution of **V-3** (1.60 g, 7.33 mmol) and KOH (4.14 g, 73.3 mmol) in THF/EtOH (200 : 100 mL) was stirred under reflux for 2.5 h. Then the solvent was removed under reduced pressure, resublimed in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O and acidified with HCl (aq). The organic layer was dried over MgSO<sub>4</sub> and the solvent was removed under reduced pressure yielded **V-4** (1.44 g, 7.1 mmol, 96%) as a white solid. IR (neat):  $\nu = 3301$  (C $\equiv$ CH), 2116 (C $\equiv$ C), 1685 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 8.06$  (d,  $^3J = 8$  Hz, 2 H, OOCArH<sub>o</sub>), 6.95 (d,  $^3J = 8$  Hz, 2 H, OOCArH<sub>m</sub>), 4.15 (t,  $^3J = 6$  Hz, 2 H, OCH<sub>2</sub>CH<sub>2</sub>), 2.43 (dt,  $^3J = 6$  Hz &  $^2J = 2$  Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>C $\equiv$ CH), 2.04 (quint.,  $^3J = 6$  Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.99 (t,  $^4J = 2$  Hz, 1H, CH<sub>2</sub>C $\equiv$ CH) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 171.4, 163.4, 132.4, 121.6, 114.2, 83.1, 69.1, 66.4, 28.0, 15.1$  ppm.

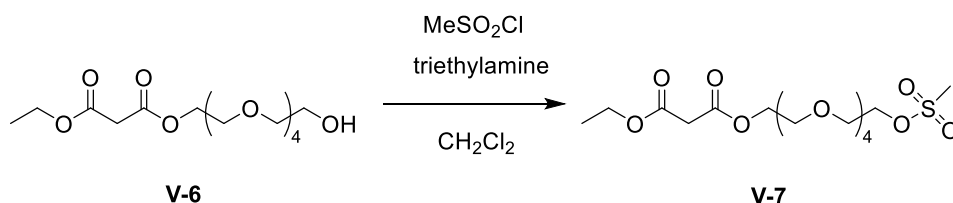




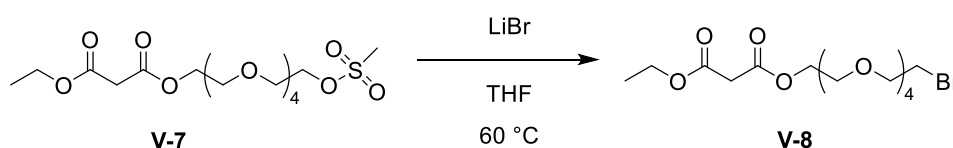
**Compound V-5:** DCC (557 mg, 2.7 mmol) was added to a solution of **V-4** (500 mg, 2.4 mmol), 5-trimethylsilyl-1-pentyn-3-ol (0.44 mL, 2.4 mmol), DMAP (61 mg, 0.5 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (40 mL) at 0°C. The resulting mixture was stirred at room temperature for 2.5 d. Then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 5:5) yielded **V-5** (546 mg, 1.6 mmol, 65%) as a colorless oil. IR (neat):  $\nu = 3301$  (C≡C-H), 2176 (C≡CSi), 2119 (C≡CH), 1713 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.98$  (d, <sup>3</sup>J = 9 Hz, 2 H, ArH<sub>o</sub>), 6.91 (d, <sup>3</sup>J = 9 Hz, 2 H, ArH<sub>m</sub>), 4.37 (t, <sup>3</sup>J = 6 Hz, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>), 4.13 (t, <sup>3</sup>J = 6 Hz, 2 H, ArOCH<sub>2</sub>CH<sub>2</sub>), 2.42 (m, 4H, CH<sub>2</sub>CH<sub>2</sub>C≡C), 2.05-1.96 (m, 5 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub> & C≡CH), 0.15 (s, 9 H, Si(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.2, 162.6, 131.5, 122.6, 114.0, 105.8, 85.2, 83.1, 69.0, 66.2, 63.2, 27.9$  (two peaks), 16.7, 15.0, 0.0 ppm.



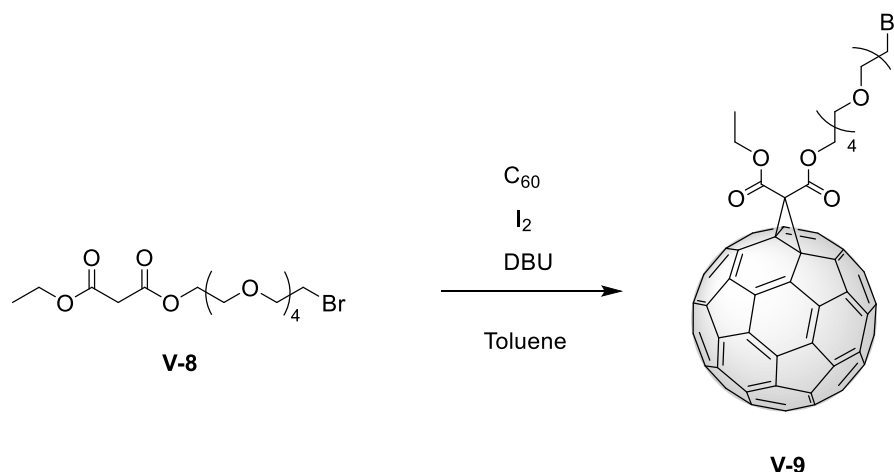
**Compound V-6:** Malonyl chloride (0.25 mL, 1.95 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (40 mL) was added dropwise to a solution of pentaerythritol (0.82 mL, 3.90 mmol) and pyridine (0.17 mL, 2.15 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (30 mL). The resulting mixture was stirred for 2 h at room temperature. The reaction mixture was filtered on SiO<sub>2</sub> and concentrated. Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/ether/MeOH 100:10:0 to 100:10:3) gave **V-6** (444 mg, 65%) as a colorless oil. IR (neat):  $\nu = 3456$  (br, OH), 1730 (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.33$  (t, <sup>3</sup>J = 4 Hz, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 4.22 (q, <sup>3</sup>J = 7 Hz, 2 H, COOCH<sub>2</sub>CH<sub>3</sub>), 3.74 (t, <sup>3</sup>J = 4 Hz, 4 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 3.71-3.66 (m, 12 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 3.43 (s, 2 H, OOCCH<sub>2</sub>COO), 2.08 (s, 1 H, OH), 1.30 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.7, 166.5, 72.5, 70.6, 70.5, 70.4, 68.8, 64.6, 61.8, 61.6, 41.5, 14.1$  ppm.



**Compound V-7:** Methanesulfonyl chloride (0.18 mL, 2.27 mmol) was added to a solution of **V-6** (400 mg, 1.14 mmol) and triethylamine (0.32 mL, 2.27 mmol) in CH<sub>2</sub>Cl<sub>2</sub> (20 mL). The resulting mixture was stirred for 1 h at room temperature. The reaction mixture was filtered on SiO<sub>2</sub> and concentrated. Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/ether/MeOH 100:7:0 to 100:10:2) gave **V-7** (462 mg, 91%) as a colorless oil. IR (neat):  $\nu = 1730$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.38$  (t, <sup>3</sup>J = 4 Hz, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 4.30 (t, <sup>3</sup>J = 5 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>OSO<sub>2</sub>CH<sub>3</sub>), 4.20 (q, <sup>3</sup>J = 7 Hz, 2 H, COOCH<sub>2</sub>CH<sub>3</sub>), 3.76 (t, <sup>3</sup>J = 4 Hz, 2 H, OCH<sub>2</sub>CH<sub>2</sub>OSO<sub>2</sub>CH<sub>3</sub>), 3.71 (t, <sup>3</sup>J = 4 Hz, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.68-3.62 (m, 12 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 3.40 (s, 2 H, OOCCH<sub>2</sub>COO), 3.08 (s, 3 H, CH<sub>2</sub>OSO<sub>2</sub>CH<sub>3</sub>), 1.27 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.7$ , 166.5, 70.6, 70.6 (four peaks), 70.5, 69.3, 69.0, 68.9, 64.5, 61.6, 41.5, 37.7, 14.1 ppm.

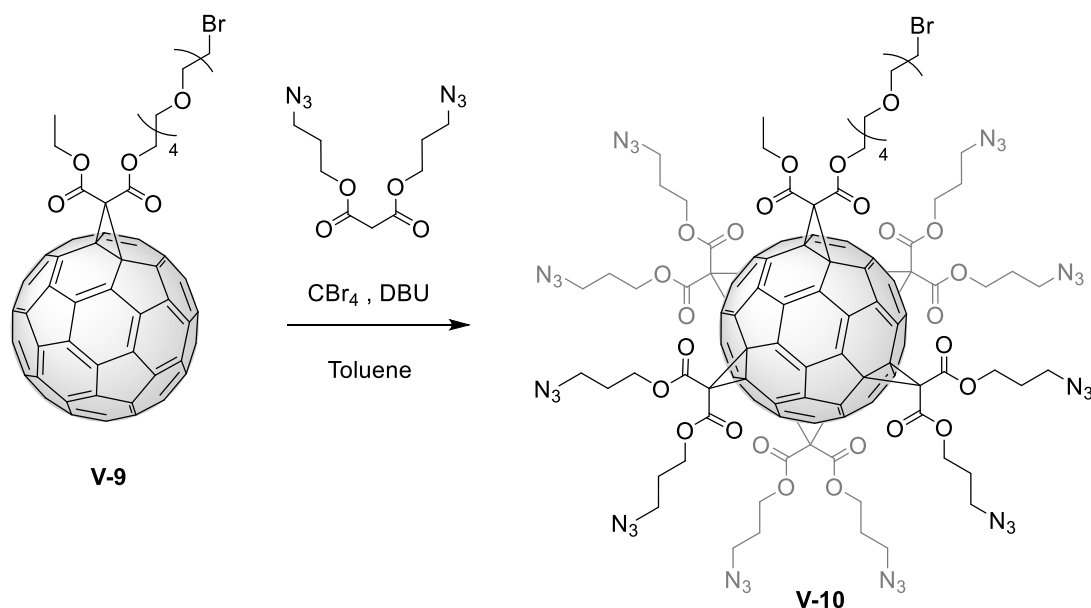


**Compound V-8:** A mixture of **V-7** (458 mg, 1.06 mmol) and LiBr (185 mg, 2.28 mmol) in THF anhydride was stirred at 60°C for 4 h. The reaction mixture was filtered on SiO<sub>2</sub> and concentrated. Column chromatography (SiO<sub>2</sub>, CH<sub>2</sub>Cl<sub>2</sub>/ether/MeOH 100:10:0 to 100:10:1) gave **V-8** (386 mg, 87%) as a colorless oil. IR (neat):  $\nu = 1731$  (C=O) cm<sup>-1</sup>. <sup>1</sup>H-NMR (400 MHz, CDCl<sub>3</sub>):  $\delta = 4.30$  (t, <sup>3</sup>J = 5 Hz, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 4.20 (q, <sup>3</sup>J = 7 Hz, 2 H, COOCH<sub>2</sub>CH<sub>3</sub>), 3.81 (t, <sup>3</sup>J = 6 Hz, 2 H, OCH<sub>2</sub>CH<sub>2</sub>Br), 3.71 (t, <sup>3</sup>J = 5 Hz, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.68-3.63 (m, 12 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 3.47 (t, <sup>3</sup>J = 6 Hz, 2 H, OCH<sub>2</sub>CH<sub>2</sub>Br), 3.40 (s, 2 H, OOCCH<sub>2</sub>COO), 1.28 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C-NMR (100 MHz, CDCl<sub>3</sub>):  $\delta = 166.7$ , 166.5, 71.2, 70.7 (two peaks), 70.6 (four peaks), 68.9, 64.6, 61.6, 41.5, 30.0, 14.1 ppm.

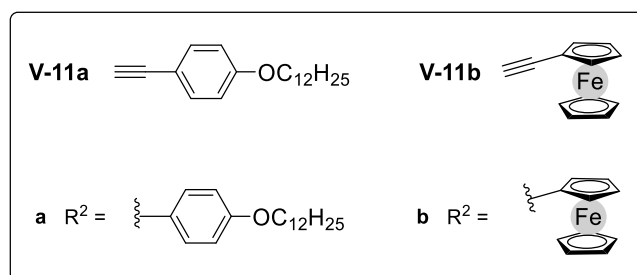
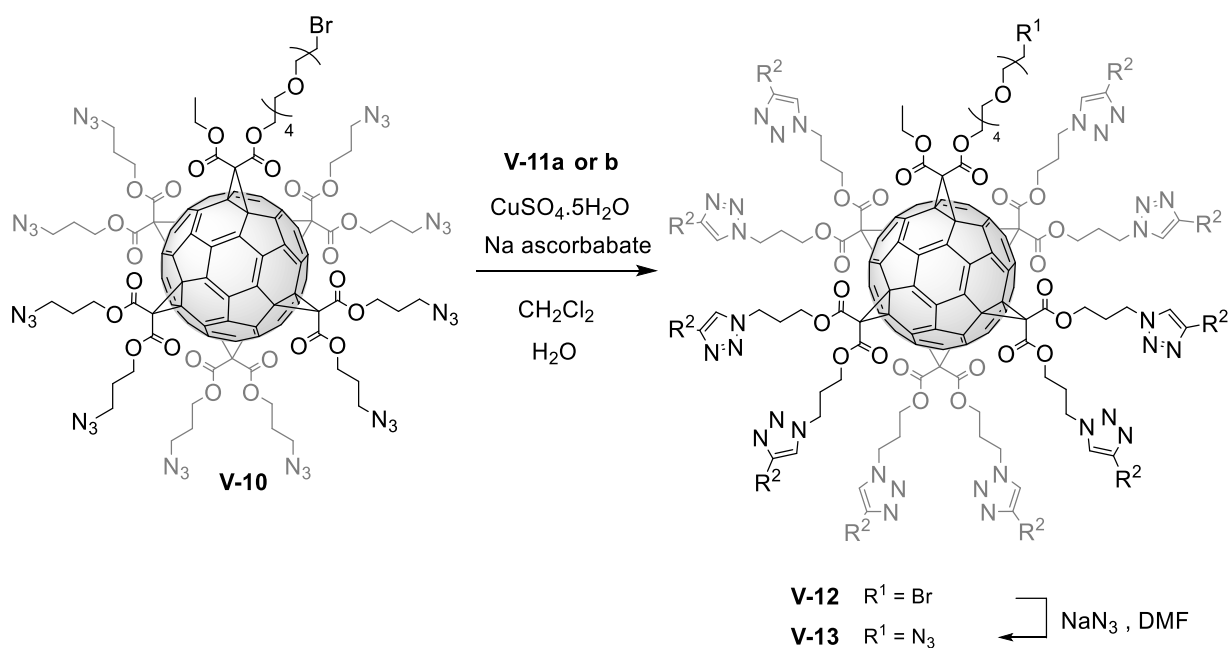


**Compound V-9:** DBU (0.37 mL, 2.5 mmol) was added to a solution of **V-8** (445 mg, 1.1 mmol), C<sub>60</sub> (850 mg, 1.2 mmol) and I<sub>2</sub> (326 mg, 1.3 mmol) in toluene (1.7 L). The resulting mixture was stirred at room temperature for 30 min, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/THF, 9:1) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/THF, 94:6) yielded **V-9** (700

mg, 0.62 mmol, 57%) as a brown glassy solid. IR (neat):  $\nu = 1743$  (C=O)  $\text{cm}^{-1}$ . UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 260$  ( $1.3 \times 10^5$ ), 330 ( $4 \times 10^4$ ), 394 (sh,  $5 \times 10^4$ ), 402 (sh,  $4 \times 10^3$ ), 414 (sh,  $3 \times 10^3$ ), 426 ( $3 \times 10^3$ ), 498 (sh,  $2 \times 10^3$ ), 631 (sh,  $3 \times 10^2$ ), 693 (sh,  $2 \times 10^2$ ) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.65$  (m, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 4.56 (q, <sup>3</sup>J = 7 Hz, 2 H, COOCH<sub>2</sub>CH<sub>3</sub>), 3.88 (m, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.81 (t, <sup>3</sup>J = 6 Hz, 2 H, OCH<sub>2</sub>CH<sub>2</sub>Br), 3.71-3.64 (m, 12 H, (OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>O), 3.47 (t, <sup>3</sup>J = 6 Hz, 2 H, OCH<sub>2</sub>CH<sub>2</sub>Br), 1.49 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.6$ , 163.4, 145.4, 145.3, 145.2 (two peaks), 144.9, 144.7, 144.6, 143.9 (two peaks), 143.1, 143.0 (three peaks), 142.2 (two peaks), 141.9 (two peaks), 141.0, 140.9, 139.2, 138.9, 71.5, 71.2, 70.7 (three peaks), 70.6 (two peaks), 68.8, 66.2, 63.5, 52.1, 30.3, 14.3 ppm. MS-MALDI-TOF: 1134.1 ([M]<sup>+</sup>, calcd. for C<sub>75</sub>H<sub>25</sub>O<sub>8</sub>Br : 1134.1).



**Compound V-10:** DBU (0.76 mL, 5.0 mmol) was added to a solution of **V-9** (285 mg, 251  $\mu\text{mol}$ ), **IV-27** (441 mg, 1.63 mmol) and CBr<sub>4</sub> (5.42 g, 16.3 mmol) in toluene (90 mL). The resulting mixture was stirred overnight at room temperature, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/Et<sub>2</sub>O, 7:3) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/Et<sub>2</sub>O, 9:1) yielded **V-10** (238 mg, 96  $\mu\text{mol}$ , 38%) as an orange glassy solid. IR (neat):  $\nu = 2095$  (N<sub>3</sub>), 1742 (C=O)  $\text{cm}^{-1}$ . <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 4.37$  (m, 24 H, COOCH<sub>2</sub>), 3.81 (t, <sup>3</sup>J = Hz, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.74-3.60 (m, 14 H, OCH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>), 3.47 (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>Br), 3.40 (t, <sup>3</sup>J = Hz, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 1.97 (m, 20 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 1.34 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>2</sub>CH<sub>3</sub>) ppm.



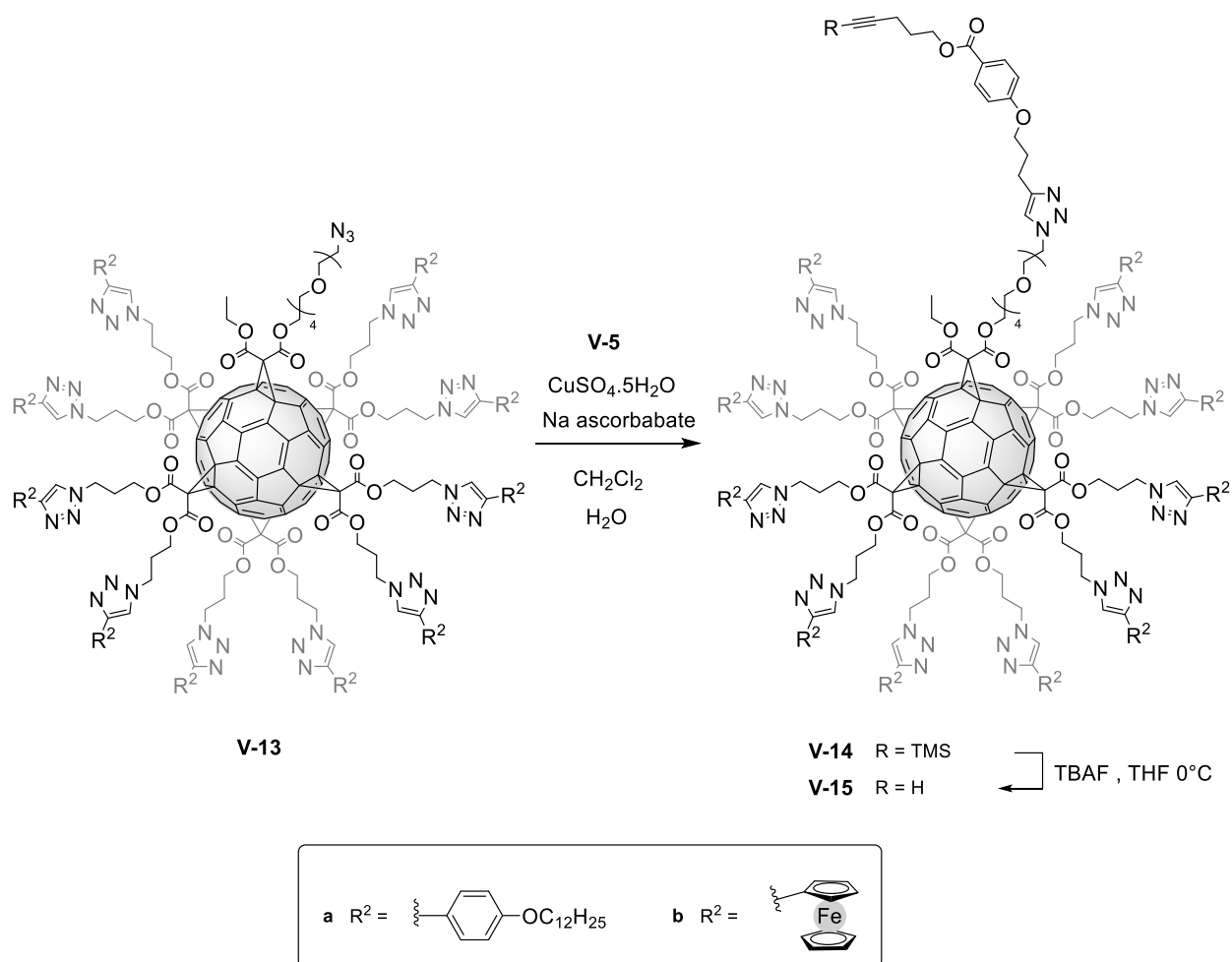
**Compound V-12a:** A solution of **V-10** (352 mg, 142  $\mu$ mol), **V-11a** (541 mg, 1.89 mmol), CuSO<sub>4</sub>·5H<sub>2</sub>O (3 mg, 11  $\mu$ mol) and sodium ascorbate (4 mg, 22  $\mu$ mol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (2 : 1 mL) was stirred overnight at 30°C, then CH<sub>2</sub>Cl<sub>2</sub> was added, washed with water and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 96:4) yielded **FS305** (737 mg, 138  $\mu$ mol, 97%) as an orange glassy solid. IR (neat):  $\nu = 1741$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 254$  (3.2x10<sup>5</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.76$ -7.65 (m, 30 H, *H*<sub>triazole</sub> & Ar*H*<sub>o</sub>), 6.87 (m, 20 H, Ar*H*<sub>m</sub>), 4.40 (m, 44 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub> & COOCH<sub>2</sub>), 3.92 (m, 20 H, OCH<sub>2</sub>(CH<sub>2</sub>)<sub>10</sub>CH<sub>3</sub>) 3.76 (t, <sup>3</sup>*J* = 6 Hz, 2 H, OCH<sub>2</sub>CH<sub>2</sub>Br), 3.60 (m, 16 H, COOCH<sub>2</sub>CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>CH<sub>2</sub>Br), 3.43 (t, <sup>3</sup>*J* = 6 Hz, 2 H, OCH<sub>2</sub>CH<sub>2</sub>Br), 2.32 (m, 20 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 1.76 (m, 20 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub>) 1.43 (m, 20 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub>), 1.26 (m, 163 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub> & COOCH<sub>2</sub>CH<sub>3</sub>), 0.87 (t, <sup>3</sup>*J* = 7 Hz, 30 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.6, 159.2, 147.8, 145.9, 141.2, 127.0, 122.9, 119.2, 114.8, 71.2, 70.5, 69.2, 68.1, 63.8, 46.8, 45.4, 31.9, 30.5, 29.7, 29.5, 29.4, 29.3, 26.1, 22.7, 14.2$  ppm. MS-MALDI-TOF: 5334.8 ([M]<sup>+</sup>, calcd for C<sub>320</sub>H<sub>385</sub>O<sub>38</sub>N<sub>30</sub>Br: 5334.8 (monoisotopic)).

**Compound V-12b:** A solution of **V-10** (238 mg, 96  $\mu$ mol), ethynylferrocene (**V-11b**) (262 mg, 1.25 mmol), CuSO<sub>4</sub>·5H<sub>2</sub>O (3 mg, 11  $\mu$ mol) and sodium ascorbate (4 mg, 22  $\mu$ mol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O

(2 : 1 mL) was stirred overnight at 30°C, then CH<sub>2</sub>Cl<sub>2</sub> was added, washed with a solution of EDTA (aq) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 95:5) yielded **V-12b** (383 mg, 84 μmol, 87%) as an orange glassy solid. IR (neat):  $\nu = 1740$  (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 232$  (2.5x10<sup>5</sup>), 269 (1.5x10<sup>5</sup>), 344 (sh, 3x10<sup>4</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.56$  (m, 10 H, *H*<sub>triazole</sub>), 4.72 (s, 20 H, *H*<sub>ferrocene</sub>), 4.39 (m, 44 H, COOCH<sub>2</sub> & CH<sub>2</sub>N<sub>triazole</sub>), 4.28 (s, 20 H, *H*<sub>ferrocene</sub>), 4.06 (s, 50 H, *H*<sub>ferrocene</sub>), 3.78 (t, <sup>3</sup>J = 6 Hz, COOCH<sub>2</sub>CH<sub>2</sub>O), 4.71-4.60 (m, 14 H, OCH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>), 3.45 (t, <sup>3</sup>J = 6 Hz, CH<sub>2</sub>CH<sub>2</sub>Br), 2.34 (m, 20 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 1.31 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.5$ , 147.0, 146.0, 145.9, 145.7, 141.5, 141.2, 119.3, 75.5, 71.2, 70.6, 69.1, 68.7, 66.7, 63.8, 46.7, 45.3, 30.5, 29.3, 14.1 ppm. MS-MALDI-TOF: 4575.1 (100%, [M]<sup>+</sup>, calcd for C<sub>240</sub>H<sub>185</sub>O<sub>28</sub>N<sub>30</sub>BrFe<sub>10</sub>: 4575.7), 3887.5 (54%, [M-(Malonate-Fe)]<sup>+</sup>, calcd for C<sub>207</sub>H<sub>153</sub>O<sub>24</sub>N<sub>24</sub>BrFe<sub>8</sub>: 3887.5).

**Compound V-13a:** **V-12a** (706 mg, 132 μmol) and NaN<sub>3</sub> (50 mg, 769 μmol) in DMF/THF (6:2 mL) were stirred at room temperature for 18 h. THF was removed under reduced pressure, then extracted with CH<sub>2</sub>Cl<sub>2</sub>, washed with H<sub>2</sub>O and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 96:4) yielded **V-13a** (650 mg, 123 μmol, 93%) as an orange glassy solid. IR (neat):  $\nu = 2101$  (N<sub>3</sub>), 1741 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.76$ -7.65 (m, 30 H, *H*<sub>triazole</sub> & Ar*H*<sub>o</sub>), 6.87 (m, 20 H, Ar*H*<sub>m</sub>), 4.40 (m, 44 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub> & COOCH<sub>2</sub>), 3.92 (m, 20 H, OCH<sub>2</sub>(CH<sub>2</sub>)<sub>10</sub>CH<sub>3</sub>), 3.60 (m, 18 H, COOCH<sub>2</sub>CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 3.43 (t, <sup>3</sup>J = 6 Hz, 2 H, OCH<sub>2</sub>CH<sub>2</sub>Br), 2.32 (m, 20 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 1.76 (m, 20 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub>) 1.43 (m, 20 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub>), 1.26 (m, 163 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub> & COOCH<sub>2</sub>CH<sub>3</sub>), 0.87 (t, <sup>3</sup>J = 7 Hz, 30 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>) ppm.

**Compound V-13b:** **V-12b** (129 mg, 28 μmol) and NaN<sub>3</sub> (11 mg, 169 μmol) in DMF (2 mL) were stirred at room temperature for 18 h. The product was precipitated with Et<sub>2</sub>O/H<sub>2</sub>O then resolubilized with CH<sub>2</sub>Cl<sub>2</sub>, washed with H<sub>2</sub>O and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 96:4) yielded **V-13b** (120 mg, 26 μmol, 94%) as an orange glassy solid. IR (neat):  $\nu = 2102$  (N<sub>3</sub>), 1743 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.56$  (m, 10 H, *H*<sub>triazole</sub>), 4.72 (s, 20 H, *H*<sub>ferrocene</sub>), 4.39 (m, 44 H, COOCH<sub>2</sub> & CH<sub>2</sub>N<sub>triazole</sub>), 4.28 (s, 20 H, *H*<sub>ferrocene</sub>), 4.06 (s, 50 H, *H*<sub>ferrocene</sub>), 3.78 (t, <sup>3</sup>J = 6 Hz, COOCH<sub>2</sub>CH<sub>2</sub>O), 4.71-4.60 (m, 14 H, OCH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>), 3.36 (t, <sup>3</sup>J = 6 Hz, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 2.34 (m, 20 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 1.31 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>2</sub>CH<sub>3</sub>) ppm.



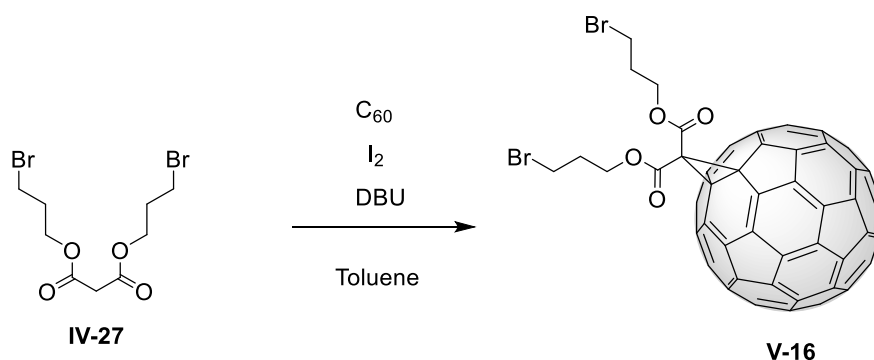
**Compound V-14a:** **V-13a** (200 mg, 38  $\mu\text{mol}$ ) was engaged in a solution of **V-5** (25 mg, 75  $\mu\text{mol}$ ),  $\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$  (1 mg, 5  $\mu\text{mol}$ ) and sodium ascorbate (3 mg, 15  $\mu\text{mol}$ ) in  $\text{CH}_2\text{Cl}_2/\text{H}_2\text{O}$  (2/1 mL). The resulting mixture was stirred at 30°C for 2.5 h, then  $\text{CH}_2\text{Cl}_2$  was added to the mixture, washed with  $\text{H}_2\text{O}$  and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 97:3) yielded **V-14a** (211 mg, 37  $\mu\text{mol}$ , 99%) as an orange glassy solid. IR (neat):  $\nu = 2178$  ( $\text{C}\equiv\text{CSi}$ ),  $1741$  ( $\text{C}=\text{O}$ )  $\text{cm}^{-1}$ . UV/Vis ( $\text{CH}_2\text{Cl}_2$ ):  $\lambda_{\text{max}}(\epsilon) = 255$  ( $2.8 \times 10^5$ ) nm.  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 7.96$  (d,  $^3J = 9$  Hz, 2 H,  $\text{OOCArH}_o$ ), 7.76-7.64 (m, 30 H,  $\text{OArH}_m$  &  $H_{\text{triazole-ArOC}_{12}}$ ), 7.47 (s, 1 H,  $H_{\text{triazole-ArCOO}}$ ), 6.87 (m, 22 H,  $\text{OOCArH}_m$  &  $\text{OArH}_o$ ), 4.46-4.27 (m, 48 H), 4.05 (t,  $^3J = 6$  Hz, 2 H,  $\text{OOCArOCH}_2\text{CH}_2$ ), 3.92 (m, 20 H,  $\text{ArOCH}_2\text{CH}_2$ ), 3.81 (t,  $^3J = 6$  Hz, 2 H,  $\text{N}_{\text{triazole}}\text{CH}_2\text{CH}_2\text{O}$ ), 3.66 (m, 2 H,  $\text{COOCH}_2\text{CH}_2\text{O}$ ), 3.57 (m, 12 H,  $\text{CH}_2(\text{OCH}_2\text{CH}_2)_3\text{OCH}_2$ ), 2.89 (t,  $^3J = 6$  Hz, 2 H,  $\text{C}_{\text{triazole}}\text{CH}_2\text{CH}_2$ ), 2.40 (t,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{C}\equiv\text{C}$ ), 2.32 (m, 20 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{N}_{\text{triazole}}$ ), 2.18 (m, 2 H,  $\text{OCH}_2\text{CH}_2\text{CH}_2\text{C}_{\text{triazole}}$ ), 1.96 (quint.,  $^3J = 6$  Hz, 2 H,  $\text{CH}_2\text{CH}_2\text{CH}_2\text{C}\equiv\text{C}$ ), 1.75 (m, 20 H,  $\text{OCH}_2\text{CH}_2\text{CH}_2$ ), 1.43 (m, 20 H,  $\text{OCH}_2\text{CH}_2\text{CH}_2\text{CH}_2$ ), 1.26 (br s, 163 H,  $\text{CH}_2(\text{CH}_2)_8\text{CH}_3$  &  $\text{COOCH}_2\text{CH}_3$ ), 0.87 (t,  $^3J = 7$  Hz, 30 H,  $\text{CH}_2\text{CH}_2\text{CH}_3$ ), 0.14 (s, 9 H,  $\text{Si}(\text{CH}_3)_3$ ) ppm.  $^{13}\text{C}$  NMR ( $\text{CDCl}_3$ , 100 MHz):  $\delta = 166.1$ , 163.5, 163.4, 162.7, 159.1, 147.7 (two peaks), 145.8, 145.6, 141.4, 141.2, 141.1, 141.0, 131.5, 126.9, 122.8, 122.5, 122.2, 119.1 (two peaks), 119.0, 114.7, 114.0, 105.8, 85.2, 70.4, 69.4, 69.0, 68.5, 68.4, 68.0, 67.0, 63.7, 63.2, 46.7, 45.3, 31.8,

29.6, 29.5 (two peaks), 29.4, 29.3, 29.2, 27.8, 26.0, 22.6, 16.7, 14.0, 0.0 ppm. MS-MALDI-TOF: 5638.5 ([M]<sup>+</sup>, calcd for C<sub>340</sub>H<sub>411</sub>O<sub>41</sub>N<sub>33</sub>Si: 5640.5 (exact mass)).

**Compound V-14b:** **V-13b** (76 mg, 17 μmol) was engaged in a solution of **V-5** (12 mg, 33 μmol), CuSO<sub>4</sub>·5H<sub>2</sub>O (1 mg, 5 μmol) and sodium ascorbate (3 mg, 15 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (2/1 mL). The resulting mixture was stirred at 30°C for 2.5 h, then CH<sub>2</sub>Cl<sub>2</sub> was added to the mixture, washed with H<sub>2</sub>O and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 96:4) yielded **V-14b** (72 mg, 15 μmol, 88%) as an orange glassy solid. IR (neat):  $\nu = 2175$  (C≡CSi), 1742 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 231$  (2.7×10<sup>5</sup>), 246 (sh, 2.1×10<sup>5</sup>), 269 (1.7×10<sup>5</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.96$  (d, <sup>3</sup>J = 9 Hz, 2 H, OOCArH<sub>o</sub>), 7.58-7.56 (m, 10 H, H<sub>triazole</sub>-Fe), 7.47 (s, 1 H, H<sub>triazole</sub>-ArCOO), 6.89 (d, <sup>3</sup>J = 9 Hz, 2 H, OOCArH<sub>m</sub>), 4.71 (br s, 20 H, H<sub>ferrocene</sub>), 4.46-4.30 (m, 48 H, COOCH<sub>2</sub> & CH<sub>2</sub>N<sub>triazole</sub>), 4.26 (br s, 20 H, H<sub>ferrocene</sub>), 4.04 (m, 52 H, H<sub>ferrocene</sub> & CH<sub>2</sub>CH<sub>2</sub>OAr), 3.81 (m, 2 H, OCH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 3.68 (m, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.56 (m, 12 H, O(CH<sub>2</sub>CH<sub>2</sub>O)<sub>3</sub>), 2.89 (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>), 2.40-2.33 (m, 22 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub> & CH<sub>2</sub>C≡C), 2.18 (m, 2 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>), 1.96 (quint., <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C≡C), 1.29 (t, <sup>3</sup>J = 7 Hz, 3 H, CH<sub>2</sub>CH<sub>3</sub>), 0.14 (s, 9 H, Si(CH<sub>3</sub>)<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.1, 163.4, 163.3, 163.2, 162.6, 146.8, 146.7, 145.8$  (two peaks), 145.7, 145.6, 141.3, 141.3, 141.0 (two peaks), 131.5, 122.5, 122.0, 119.2, 114.0, 105.8, 85.2, 75.2, 70.5, 70.4, 70.3, 69.5, 69.0, 68.6, 67.0, 66.6, 63.7, 63.2, 50.0, 46.6, 45.2, 29.2, 28.7, 27.8, 22.0, 16.7, 13.9, 0.0 ppm. MS-MALDI-TOF: 4879.2 ([M]<sup>+</sup>, calcd for C<sub>260</sub>H<sub>211</sub>O<sub>31</sub>N<sub>33</sub>SiFe<sub>10</sub> : 4880.2 (molecular weight)).

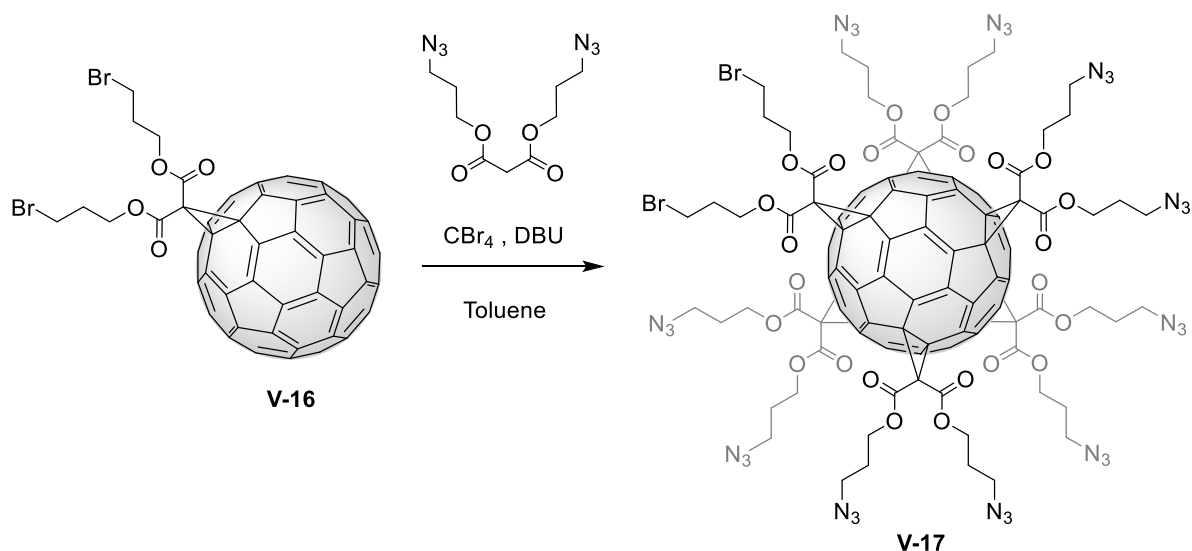
**Compound V-15a:** TBAF 1M in THF (0.24 mL, 240 μmol) was added to a solution of **V-14a** (903 mg, 160 μmol) in THF (15 mL) at 0°C. The resulting mixture was stirred at 0°C for 20 min, then water was added. THF was removed under reduced pressure and the product was solubilized in CH<sub>2</sub>Cl<sub>2</sub>. The organic layer was washed with water and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 97:3) yielded **V-15a** (884 mg, 159 μmol, 99%) as an orange glassy solid. IR (neat):  $\nu = 3315$  (C≡C-H), 1741 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon) = 256$  (2.6×10<sup>5</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.96$  (d, <sup>3</sup>J = 9 Hz, 2 H, OOCArH<sub>o</sub>), 7.76-7.64 (m, 30 H, OArH<sub>m</sub> & H<sub>triazole</sub>-ArOC<sub>12</sub>), 7.47 (s, 1 H, H<sub>triazole</sub>-ArCOO), 6.94 (d, <sup>3</sup>J = 9 Hz, 2 H, OOCArH<sub>m</sub>), 6.87 (m, 20 H, OArH<sub>o</sub>), 4.55 (t, <sup>3</sup>J = 6 Hz, 2 H, ArCOOCH<sub>2</sub>CH<sub>2</sub>), 4.46-4.27 (m, 48 H), 4.05 (t, <sup>3</sup>J = 6 Hz, 2 H, OOCArOCH<sub>2</sub>CH<sub>2</sub>), 3.92 (m, 20 H, ArOCH<sub>2</sub>CH<sub>2</sub>), 3.81 (t, <sup>3</sup>J = 6 Hz, 2 H, N<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>O), 3.67 (m, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.57 (m, 12 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 2.89 (t, <sup>3</sup>J = 6 Hz, 2 H, C<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>), 2.37-2.27 (m, 22 H, CH<sub>2</sub>CH<sub>2</sub>C≡C & COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 2.17 (m, 2 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>), 1.97 (m, 3 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C≡CH), 1.75 (m, 20 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.43 (m, 20 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.26 (br s, 163 H, CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub> & COOCH<sub>2</sub>CH<sub>3</sub>), 0.87 (t, <sup>3</sup>J = 7 Hz, 30 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.2, 163.5, 162.8, 159.2, 147.8$  (two peaks), 145.9, 141.2, 131.6, 127.0, 123.0, 122.5, 122.1, 119.2, 114.8, 114.1, 83.1, 70.5, 69.5, 69.1 (two peaks), 68.1, 67.1, 63.8, 63.2, 56.8, 45.4, 31.9, 29.7 (two peaks), 29.6, 29.5, 29.4, 29.3, 28.8, 27.8, 26.1, 22.7, 22.1, 15.4, 14.1 ppm. MS-MALDI-TOF: 5566.3 ([M]<sup>+</sup>, calcd for C<sub>337</sub>H<sub>403</sub>O<sub>41</sub>N<sub>33</sub>: 5568.0 (exact mass)).

**Compound V-15b:** TBAF 1M in THF (0.02 mL, 18  $\mu$ mol) was added to a solution of **V-14b** (44 mg, 9  $\mu$ mol) in THF (4 mL) at 0°C. The resulting mixture was stirred at 0°C for 20 min, then water was added. THF was removed under reduced pressure and the product was solubized in CH<sub>2</sub>Cl<sub>2</sub>. The organic layer was washed with water and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 96:4) yielded **V-15b** (40 mg, 8.3  $\mu$ mol, 92%) as an orange glassy solid. IR (neat):  $\nu$  = 3296 (C $\equiv$ C-H), 1741 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\max}(\epsilon)$  = 269 (sh, 9x10<sup>4</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  = 7.96 (d, <sup>3</sup>J = 9 Hz, 2 H, OOCArH<sub>o</sub>), 7.58-7.56 (m, 10 H, H<sub>triazole</sub>-Fe), 7.47 (s, 1 H, H<sub>triazole</sub>-ArCOO), 6.89 (d, <sup>3</sup>J = 9 Hz, 2 H, OOCArH<sub>m</sub>), 4.72 (br s, 20 H, H<sub>ferrocene</sub>), 4.46-4.30 (m, 48 H, COOCH<sub>2</sub> & CH<sub>2</sub>N<sub>triazole</sub>), 4.28 (br s, 20 H, H<sub>ferrocene</sub>), 4.06 (m, 52 H, H<sub>ferrocene</sub> & CH<sub>2</sub>CH<sub>2</sub>OAr), 3.82 (m, 2 H, OCH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 3.68 (m, 2 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.57 (m, 12 H, O(CH<sub>2</sub>CH<sub>2</sub>O)<sub>3</sub>), 2.89 (t, <sup>3</sup>J = 6 Hz, 2 H, CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>), 2.40-2.33 (m, 22 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub> & CH<sub>2</sub>C $\equiv$ C), 2.18 (m, 2 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>), 1.98 (m, 3 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C $\equiv$ CH), 1.30 (m, 3 H, CH<sub>2</sub>CH<sub>3</sub>), ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  = 166.3, 163.5, 162.8, 147.0, 146.8, 145.9, 141.1, 131.6, 122.5, 122.1, 119.3, 114.1, 83.2, 75.4, 70.5, 69.6, 69.1, 68.7, 67.1, 66.7, 63.8, 63.2, 50.1, 46.7, 45.1, 29.3, 28.8, 27.8, 22.1, 15.4, 14.1 ppm. MS-MALDI-TOF : 4808.1 ([M]<sup>+</sup>, calcd for C<sub>257</sub>H<sub>203</sub>O<sub>31</sub>N<sub>33</sub>Fe<sub>10</sub>: 4808.0 (molecular weight)).

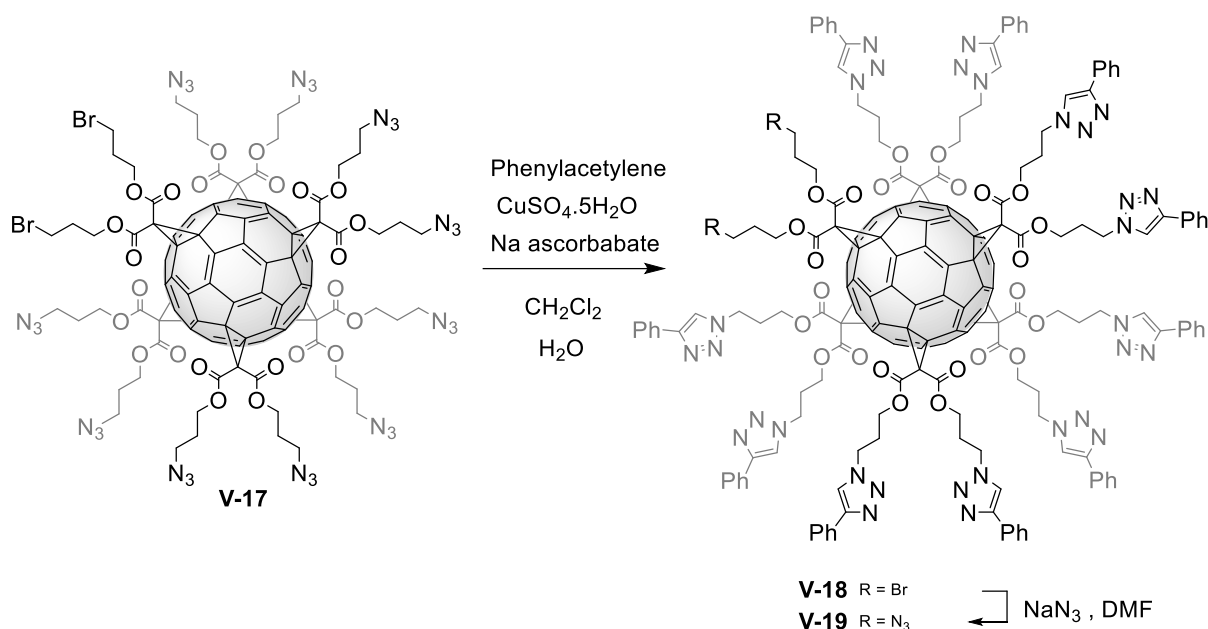


**Compound V-16:** DBU (0.18 mL, 1.2 mmol) was added to a solution of **IV-27** (191 mg, 0.55 mmol), C<sub>60</sub> (400 mg, 0.55 mmol) and I<sub>2</sub> (169 mg, 0.66 mmol) in toluene (0.8 L). The resulting mixture was stirred at room temperature for 30 min, then filtered on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>/cyclohexane, 3:7) yielded **V-16** (302 mg, 0.28 mmol, 51%) as a brown glassy solid. IR (neat):  $\nu$  = 1743 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  = 4.67 (t, <sup>3</sup>J = 6 Hz, 4 H, COOCH<sub>2</sub>CH<sub>2</sub>), 3.58 (t, <sup>3</sup>J = 6 Hz, 4 H, CH<sub>2</sub>CH<sub>2</sub>Br), 2.41 (quint., <sup>3</sup>J = 6 Hz, 4 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  = 163.4, 145.3, 145.2, 145.0, 144.9, 144.7 (two peaks), 144.6, 143.9, 143.1 (two peaks), 143.0, 142.2, 141.9, 141.0, 139.0, 71.3, 65.0, 51.8, 31.4, 29.0 ppm.





**Compound V-17:** DBU (0.23 mL, 1.3 mmol) was added to a solution of **V-16** (80 mg, 75  $\mu$ mol), **IV-12** (142 mg, 0.53 mmol) and  $\text{CBr}_4$  (1.74 g, 5.3 mmol) in toluene (50 mL). The resulting mixture was stirred overnight at room temperature, then filtered on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2/\text{Et}_2\text{O}$ , 7:3) and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{Et}_2\text{O}$ , 9:1) yielded **V-17** (100 mg, 42  $\mu$ mol, 55%) as an orange glassy solid. IR (neat):  $\nu = 2094$  ( $\text{N}_3$ ),  $1740$  ( $\text{C}=\text{O}$ )  $\text{cm}^{-1}$ .  $^1\text{H}$  NMR ( $\text{CDCl}_3$ , 400 MHz):  $\delta = 4.44$  (t,  $^3J = 6$  Hz, 4 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{Br}$ ),  $4.37$  (t,  $^3J = 6$  Hz, 20 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{N}_3$ ),  $3.47$  (t,  $^3J = 6$  Hz, 4 H,  $\text{CH}_2\text{CH}_2\text{Br}$ ),  $3.40$  (t,  $^3J = 6$  Hz, 20 H,  $\text{CH}_2\text{CH}_2\text{N}_3$ ),  $2.25$  (quint.,  $^3J = 6$  Hz, 4 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{Br}$ ),  $1.97$  (quint.,  $^3J = 6$  Hz, 20 H,  $\text{COOCH}_2\text{CH}_2\text{CH}_2\text{N}_3$ ) ppm.



**Compound V-18:** A solution of **V-17** (100 mg, 42  $\mu$ mol), phenylacetylene (0.06 mL, 499  $\mu$ mol),  $\text{CuSO}_4 \cdot 5\text{H}_2\text{O}$  (3 mg, 11  $\mu$ mol) and sodium ascorbate (4 mg, 22  $\mu$ mol) in  $\text{CH}_2\text{Cl}_2/\text{H}_2\text{O}$  (2 : 1 mL) was stirred overnight at  $30^\circ\text{C}$ , then  $\text{CH}_2\text{Cl}_2$  was added, washed with water and concentrated. Column chromatography on  $\text{SiO}_2$  ( $\text{CH}_2\text{Cl}_2$  to  $\text{CH}_2\text{Cl}_2/\text{MeOH}$ , 96:4) yielded **V-18** (113 mg, 33

μmol, 79%) as an orange glassy solid. IR (neat):  $\nu = 1742$  (C=O)  $\text{cm}^{-1}$ . UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 246$  ( $1.7 \times 10^5$ ),  $283$  ( $6 \times 10^4$ ) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.84$ - $7.75$  (m, 30 H, ArH<sub>o</sub> & H<sub>triazole</sub>),  $7.37$ - $7.28$  (m, 30 H, ArH<sub>m,p</sub>),  $4.47$ - $4.32$  (m, 44 H, COOCH<sub>2</sub>CH<sub>2</sub> & CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>),  $3.36$  (t, <sup>3</sup>J = 6 Hz, 4 H, CH<sub>2</sub>CH<sub>2</sub>Br),  $2.37$ - $2.28$  (m, 20 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>),  $2.17$  (m, 4 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 163.5$ ,  $163.4$ ,  $163.3$ ,  $147.9$  (two peaks),  $145.9$  (three peaks),  $145.8$ ,  $141.3$ ,  $141.2$  (two peaks),  $141.1$ ,  $130.5$ ,  $128.9$ ,  $128.2$ ,  $125.7$ ,  $120.1$ ,  $69.2$  (two peaks),  $69.1$ ,  $64.9$ ,  $63.8$ ,  $63.7$  (two peaks),  $46.8$ ,  $45.4$ ,  $31.1$ ,  $29.3$ ,  $29.1$  ppm. MS-MALDI-TOF :  $3429.6$  ([M]<sup>+</sup>, calcd for C<sub>194</sub>H<sub>132</sub>O<sub>24</sub>N<sub>30</sub>Br<sub>2</sub>:  $3427.1$  (molecular mass)).

**Compound V-19:** **V-18** (42 mg, 12 μmol) and NaN<sub>3</sub> (3 mg, 37 μmol) in DMF (2 mL) were stirred at room temperature. After 17 h, the product was precipitated with H<sub>2</sub>O/Et<sub>2</sub>O and resolubilized in CH<sub>2</sub>Cl<sub>2</sub>, washed with H<sub>2</sub>O and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to CH<sub>2</sub>Cl<sub>2</sub>/MeOH, 96:4) yielded **V-19** (38 mg, 11 μmol, 93%) as an orange glassy solid. IR (neat):  $\nu = 2098$  (N<sub>3</sub>),  $1741$  (C=O)  $\text{cm}^{-1}$ . <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.84$ - $7.75$  (m, 30 H, ArH<sub>o</sub> & H<sub>triazole</sub>),  $7.38$ - $7.29$  (m, 30 H, ArH<sub>m,p</sub>),  $4.48$ - $4.29$  (m, 44 H, COOCH<sub>2</sub>CH<sub>2</sub> & CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>),  $3.31$  (t, <sup>3</sup>J = 6 Hz, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>),  $2.34$  (m, 20 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>),  $1.89$  (quint., <sup>3</sup>J = 6 Hz, 4 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>) ppm.

**Compound V-20:** A solution of **V-19** (28 mg, 8 μmol), **V-15a** (97 mg, 17 μmol), CuSO<sub>4</sub>·5H<sub>2</sub>O (3 mg, 11 μmol) and sodium ascorbate (4 mg, 22 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (2 : 1 mL) was stirred overnight at 30°C, then CH<sub>2</sub>Cl<sub>2</sub> was added, washed with water and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to THF/MeOH, 92:8) followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) yielded **V-20** (68 mg, 4.7 μmol, 56%) as an orange glassy solid. IR (neat):  $\nu = 1742$  (C=O)  $\text{cm}^{-1}$ . UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 251$  ( $8.9 \times 10^5$ ) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.92$  (d, <sup>3</sup>J = 9 Hz, 4 H, OOCArH<sub>o</sub>),  $7.86$ - $7.64$  (m, 90 H, OArH<sub>m</sub> & H<sub>triazole</sub>-ArOC<sub>12</sub> & ArH<sub>o</sub> & H<sub>triazole</sub>-Ar),  $7.47$  (m, 4 H, H<sub>triazole</sub>-ArCOO),  $7.36$ - $7.27$  (m, 30 H, ArH<sub>m,p</sub>),  $6.86$  (m, 44 H, OArH<sub>o</sub> & OOCArH<sub>m</sub>),  $4.48$ - $4.27$  (m, 144 H),  $4.02$  (m, 4 H, OOCArOCH<sub>2</sub>CH<sub>2</sub>),  $3.91$  (m, 40 H, ArOCH<sub>2</sub>CH<sub>2</sub>),  $3.80$  (m, 4 H, N<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>O),  $3.65$  (m, 4 H, COOCH<sub>2</sub>CH<sub>2</sub>O),  $3.56$  (m, 24 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>),  $2.87$ - $2.80$  (m, 8 H, C<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>),  $2.31$  (m, 60 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>),  $2.17$ - $2.08$  (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>),  $1.75$  (m, 40 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>),  $1.42$  (m, 40 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>),  $1.26$  (br s, 326 H, CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub> & COOCH<sub>2</sub>CH<sub>3</sub>)  $0.87$  (br s, 60 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.3$ ,  $163.5$ ,  $163.4$  (two peaks),  $162.8$ ,  $159.2$ ,  $147.8$ ,  $145.9$ ,  $145.7$ ,  $141.2$ ,  $131.6$ ,  $130.5$ ,  $128.9$ ,  $128.2$ ,  $127.0$ ,  $125.7$ ,  $123.0$ ,  $122.5$ ,  $122.0$ ,  $120.1$ ,  $119.2$ ,  $114.8$ ,  $114.1$ ,  $70.6$ ,  $70.5$ ,  $69.5$ ,  $69.2$ ,  $68.5$ ,  $68.1$ ,  $67.2$ ,  $66.0$ ,  $63.8$ ,  $46.8$  (two peaks),  $45.5$ ,  $45.4$ ,  $31.9$ ,  $29.7$  (two peaks),  $29.5$ ,  $29.4$ ,  $29.3$ ,  $28.8$ ,  $28.4$ ,  $26.1$ ,  $22.7$ ,  $22.1$ ,  $14.1$ ,  $14.0$  ppm. MS-MALDI-TOF:  $14487.4$  ([M]<sup>+</sup>, calcd for C<sub>868</sub>H<sub>938</sub>O<sub>106</sub>N<sub>102</sub>:  $14485.1$  (exact mass)).

**Compound V-21:** A solution of **IV-18** (22 mg, 8 μmol), **V-15a** (132 mg, 24 μmol), CuSO<sub>4</sub>·5H<sub>2</sub>O (3 mg, 11 μmol) and sodium ascorbate (4 mg, 22 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (2 : 1 mL) was stirred overnight at 30°C, then CH<sub>2</sub>Cl<sub>2</sub> was added, washed with water and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to THF/MeOH, 9:1) followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) yielded **V-21** (95 mg, 4.9 μmol, 62%) as an orange

glassy solid. IR (neat):  $\nu = 1742$  (C=O)  $\text{cm}^{-1}$ . UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 253$  ( $9.2 \times 10^5$ ) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.94$  (d, <sup>3</sup>J = 9 Hz, 6 H, OOCArH<sub>o</sub>), 7.87-7.66 (m, 110 H, OArH<sub>m</sub> & H<sub>triazole</sub>-ArOC<sub>12</sub> & ArH<sub>o</sub> & H<sub>triazole</sub>-Ar), 7.47 (m, 6 H, H<sub>triazole</sub>-ArCOO), 7.36-7.27 (m, 18 H, ArH<sub>m,p</sub>), 6.87 (m, 66 H, OArH<sub>o</sub> & OOCArH<sub>m</sub>), 4.46-4.27 (m, 186 H), 4.04 (m, 6 H, OOCArOCH<sub>2</sub>CH<sub>2</sub>), 3.91 (m, 60 H, ArOCH<sub>2</sub>CH<sub>2</sub>), 3.81 (m, 6 H, N<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>O), 3.66 (m, 6 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.57 (m, 36 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 2.86 (m, 12 H, C<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>), 2.31 (m, 78 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 2.15 (m, 12 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>), 1.75 (m, 60 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.42 (m, 60 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.26 (br s, 498 H, CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub> & COOCH<sub>2</sub>CH<sub>3</sub>) 0.87 (t, <sup>3</sup>J = 7 Hz, 90 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.2, 163.5, 163.4, 162.8, 159.2, 147.9, 147.8, 147.7, 145.9, 145.8, 141.1, 131.6, 130.5, 128.8, 128.2, 127.0, 125.7, 123.0, 122.6, 122.2, 120.1, 119.2, 114.8, 114.1, 70.5, 69.5, 69.2, 69.1, 69.0, 68.5, 68.1, 67.2, 63.8$  (two peaks), 63.7, 63.6, 63.5, 46.9, 46.8 (two peaks), 45.4 (two peaks), 31.9, 29.7 (two peaks), 29.6, 29.5, 29.4, 29.3, 26.1, 22.7, 14.1 ppm.

**Compound V-22:** A solution of **IV-24** (16 mg, 5.1  $\mu\text{mol}$ ), **V-15a** (116 mg, 21  $\mu\text{mol}$ ), CuSO<sub>4</sub>.5H<sub>2</sub>O (3 mg, 11  $\mu\text{mol}$ ) and sodium ascorbate (4 mg, 22  $\mu\text{mol}$ ) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (2 : 1 mL) was stirred overnight at 30°C, then CH<sub>2</sub>Cl<sub>2</sub> was added, washed with water and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to THF/MeOH, 9:1) followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) yielded **V-22** (88 mg, 3.4  $\mu\text{mol}$ , 68%) as an orange glassy solid. IR (neat):  $\nu = 1742$  (C=O)  $\text{cm}^{-1}$ . UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 253$  ( $1.4 \times 10^6$ ) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.94$ -7.63 (m, 160 H, OArH<sub>m</sub> & H<sub>triazole</sub>-ArOC<sub>12</sub> & ArH<sub>o</sub> & H<sub>triazole</sub>-Ar & OOCArH<sub>o</sub>), 7.47 (m, 8 H, H<sub>triazole</sub>-ArCOO), 7.36-7.27 (m, 24 H, ArH<sub>m,p</sub>), 6.87 (m, 88 H, OArH<sub>o</sub> & OOCArH<sub>m</sub>), 4.46-4.27 (m, 240 H), 4.04 (m, 8 H, OOCArOCH<sub>2</sub>CH<sub>2</sub>), 3.91 (m, 80 H, ArOCH<sub>2</sub>CH<sub>2</sub>), 3.81 (m, 8 H, N<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>O), 3.66 (m, 8 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.57 (m, 48 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 2.86 (m, 16 H, C<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>), 2.31 (m, 104 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 2.15 (m, 16 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>), 1.75 (m, 80 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.42 (m, 80 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.26 (br s, 652 H, CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub> & COOCH<sub>2</sub>CH<sub>3</sub>) 0.87 (br s, 120 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta = 166.3, 163.6, 163.5$  (two peaks), 163.4, 162.8, 159.2, 147.8, 145.9, 145.7, 141.3, 141.2, 141.1, 131.6, 130.5, 128.9, 128.2, 127.0, 125.7, 123.0, 122.5, 122.2, 120.1, 119.3, 114.8, 114.1, 70.5, 69.5, 69.2, 68.5, 68.1, 67.2, 66.0, 63.8, 50.1, 46.8, 45.6, 45.4, 31.9, 29.7 (two peaks), 29.5, 29.4, 29.3, 26.1, 22.7, 14.1 ppm.

**Compound V-23:** A solution of **IV-16** (16 mg, 6.4  $\mu\text{mol}$ ), **V-15a** (293 mg, 53  $\mu\text{mol}$ ), CuSO<sub>4</sub>.5H<sub>2</sub>O (3 mg, 11  $\mu\text{mol}$ ) and sodium ascorbate (4 mg, 22  $\mu\text{mol}$ ) in CH<sub>2</sub>Cl<sub>2</sub>/H<sub>2</sub>O (2 : 1 mL) was stirred at 30°C for 4 days, then CH<sub>2</sub>Cl<sub>2</sub> was added, washed with water and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub> to THF/MeOH, 9:1) followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) yielded **V-23** (212 mg, 4.5  $\mu\text{mol}$ , 70%) as an orange glassy solid. IR (neat):  $\nu = 1742$  (C=O)  $\text{cm}^{-1}$ . UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>):  $\lambda_{\text{max}}(\epsilon) = 255$  ( $2.8 \times 10^6$ ) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta = 7.94$  (m, 16 H, OOCArH<sub>o</sub>), 7.76-7.66 (m, 240 H, OArH<sub>m</sub> & H<sub>triazole</sub>-ArOC<sub>12</sub>), 7.46 (m, 16 H, H<sub>triazole</sub>-ArCOO), 6.87 (m, 176 H, OArH<sub>o</sub> & OOCArH<sub>m</sub>), 4.46-4.27 (m, 424 H), 4.05 (m, 16 H, OOCArOCH<sub>2</sub>CH<sub>2</sub>), 3.91 (m, 160 H, ArOCH<sub>2</sub>CH<sub>2</sub>), 3.82 (m, 16 H,

N<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>O), 3.66 (m, 16 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.56 (m, 96 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 3.38 (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 2.86 (m, 32 H, C<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>), 2.32 (m, 160 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 2.16 (m, 40 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub> & CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>Br), 1.75 (m, 160 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.42 (m, 160 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.26 (br s, 1304 H, CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub> & COOCH<sub>2</sub>CH<sub>3</sub>) 0.87 (br s, 240 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz): δ = 166.3, 163.6, 163.5 (two peaks), 163.4, 163.3, 162.8, 159.2, 147.8, 145.9, 141.2, 131.6, 127.0, 123.0, 122.6, 122.1, 119.2, 114.8, 114.1, 71.4, 70.5, 69.5, 69.1, 68.6, 68.1, 67.2, 65.3, 63.8, 50.1, 46.8, 45.4, 31.9, 31.2, 29.7 (two peaks), 29.5, 29.4, 29.3, 28.5, 26.1, 22.7, 22.3, 14.1 ppm.

**Compound V-24:** **V-23** (110 mg, 2.3 μmol) and NaN<sub>3</sub> (22 mg, 338 μmol) in DMF (2 mL) were stirred at room temperature. After 17 h, the product was precipitated with H<sub>2</sub>O/Et<sub>2</sub>O and resolubilized in CH<sub>2</sub>Cl<sub>2</sub>. The organic layer was washed with H<sub>2</sub>O, dried over MgSO<sub>4</sub> and concentrated. Gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) yielded **V-24** (97 mg, 2.1 μmol, 88%) as an orange glassy solid. IR (neat): ν = 2099 (N<sub>3</sub>), 1741 (C=O) cm<sup>-1</sup>. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): δ = 7.94 (m, 16 H, OOCArH<sub>o</sub>), 7.76-7.66 (m, 240 H, OArH<sub>m</sub> & H<sub>triazole</sub>-ArOC<sub>12</sub>), 7.47 (m, 16 H, H<sub>triazole</sub>-ArCOO), 6.87 (m, 176 H, OArH<sub>o</sub> & OOCArH<sub>m</sub>), 4.46-4.27 (m, 424 H), 4.05 (m, 16 H, OOCArOCH<sub>2</sub>CH<sub>2</sub>), 3.91 (m, 160 H, ArOCH<sub>2</sub>CH<sub>2</sub>), 3.82 (m, 16 H, N<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>O), 3.66 (m, 16 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.57 (m, 96 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 3.36 (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 2.86 (m, 32 H, C<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>), 2.32 (m, 160 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 2.16 (m, 32 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>), 1.95 (m, 8 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>3</sub>), 1.75 (m, 160 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.42 (m, 160 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.26 (br s, 1304 H, CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub> & COOCH<sub>2</sub>CH<sub>3</sub>) 0.87 (br s, 240 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>) ppm.

**Compound V-25:** A solution of **V-24** (94 mg, 2 μmol), **V-15b** (60 mg, 12 μmol), CuSO<sub>4</sub>·5H<sub>2</sub>O (3 mg, 11 μmol) and sodium ascorbate (4 mg, 22 μmol) in CH<sub>2</sub>Cl<sub>2</sub>/DMF/H<sub>2</sub>O (2 : 1 : 1 mL) was stirred at 37°C for 7 days, then CH<sub>2</sub>Cl<sub>2</sub> was added, washed with water and concentrated. Column chromatography on SiO<sub>2</sub> (CH<sub>2</sub>Cl<sub>2</sub>) followed by gel permeation chromatography (Biobeads SX-1, CH<sub>2</sub>Cl<sub>2</sub>) yielded **V-25** (95 mg, 1.4 μmol, 72%) as an orange glassy solid. IR (neat): ν = 1741 (C=O) cm<sup>-1</sup>. UV/Vis (CH<sub>2</sub>Cl<sub>2</sub>): λ<sub>max</sub>(ε) = 254 (2.7×10<sup>6</sup>) nm. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): δ = 7.93 (m, 24 H, OOCArH<sub>o</sub>), 7.76-7.59 (m, 280 H, OArH<sub>m</sub> & H<sub>triazole</sub>-ArOC<sub>12</sub> & H<sub>triazole</sub>-Fe), 7.45 (m, 24 H, H<sub>triazole</sub>-ArCOO), 6.87 (m, 184 H, OArH<sub>o</sub> & OOCArH<sub>m</sub>), 4.70 (m, 80 H, H<sub>ferrocene</sub>), 4.48-4.29 (m, 624 H, COOCH<sub>2</sub> & CH<sub>2</sub>N<sub>triazole</sub>), 4.26 (m, 80 H, H<sub>ferrocene</sub>), 4.04 (m, 224 H, H<sub>ferrocene</sub> & CH<sub>2</sub>CH<sub>2</sub>OAr), 3.91 (m, 160 H, ArOCH<sub>2</sub>CH<sub>2</sub>), 3.82 (m, 24 H, N<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>O), 3.68 (m, 24 H, COOCH<sub>2</sub>CH<sub>2</sub>O), 3.57 (m, 144 H, CH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OCH<sub>2</sub>), 2.86 (m, 48 H, C<sub>triazole</sub>CH<sub>2</sub>CH<sub>2</sub>), 2.33 (m, 264 H, COOCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>N<sub>triazole</sub>), 2.15 (m, 48 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>C<sub>triazole</sub>), 1.75 (m, 160 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.42 (m, 160 H, OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>), 1.25 (m, 1316 H, CH<sub>2</sub>(CH<sub>2</sub>)<sub>8</sub>CH<sub>3</sub> & COOCH<sub>2</sub>CH<sub>3</sub>), 0.87 (m, 240 H, CH<sub>2</sub>CH<sub>2</sub>CH<sub>3</sub>) ppm. <sup>13</sup>C NMR (CDCl<sub>3</sub>, 100 MHz): δ = 166.3, 163.5, 163.4, 162.8, 159.2, 147.8, 146.9, 145.9, 145.7, 141.3, 141.2, 141.1, 131.6, 127.0, 123.0, 122.5, 122.0, 119.2, 119.2, 114.8, 114.1, 75.4, 70.5, 69.6, 69.1, 68.7, 68.1, 67.2, 66.7, 63.8, 50.1, 46.7, 45.4, 31.9, 29.7, 29.6, 29.5, 29.4, 29.3, 28.9, 28.5, 26.1, 22.7, 22.3, 22.2, 14.1, 14.0 ppm.



## **Chapter VI : General conclusions.**

The first part of this work is concerned with the development of new methodologies for the regioselective multifunctionalization of  $C_{60}$ . Our main concern was to include sub-units allowing to easily generate useful fullerene building blocks.

In a first instance, a macrocyclic approach has been investigated. For this purpose, bis-malonate macrocycles incorporating silyl groups were synthesized. Three parameters of the macrocycles have been modified: (1) the ring size, (2) the rigidity/flexibility of the spacer unit and (3) the nature of the protecting silyl groups. The reaction of the macrocycles bis-malonate with  $C_{60}$  gave three regioisomers of  $C_{60}$  bis-adducts with good regioselectivity. The cleavage of the silyl groups has been easily performed to afford the corresponding acyclic fullerene polyols. This new strategy opens new perspectives for the post-functionalization of fullerene multi-adducts derivatives.

Macrocyclic tris-malonates have been also synthesized. A stepwise synthesis has been developed in order to form the macrocycle in sufficient amount. For two macrocyclic tris-malonates, excellent results have been obtained with the exclusive formation of the  $C_{60}$  *e,e,e* tris-adduct.

The macrocyclic approach has proved to be interesting for the regioselective functionalization of  $C_{60}$ . However, in the case of the macrocyclic tris-malonates, the high number of steps for the synthesis of the macrocycles led us to develop a “tether-directed” type approach for the regioselective tris-functionalization of  $C_{60}$ . This strategy relies on a threefold Bingel reaction between  $C_{60}$  and a trialkoxysilane derivative bearing three malonate substituents. The  $C_{60}$  tris-adducts have been obtained in only three steps from commercially available starting materials. High regioselectivity has been observed for the *e,e,e* addition pattern. The easy deprotection of the silyl group afforded  $C_{60}$  tris-adducts bearing three alcohol functions which can be post-functionalized.

Optically pure *e,e,e* fullerene tris-adducts have been also prepared by using our synthetic approach. In this case, optically pure tris-malonates have been used to functionalize the fullerene core thus affording easily separable diastereoisomers differing by the absolute configuration of the inherently chiral addition pattern on the fullerene core. The absolute configuration of the inherently chiral *e,e,e* addition pattern has been unambiguously determined using X-ray crystal structure analysis for the first time.

Our “tether-directed” strategy furnishes a fast, simple, regio- and stereoselective access to  $C_{60}$  *e,e,e* tris-adduct suitable for the formation of  $C_{60}$  hexa-adducts with an octahedral addition pattern.

The second part of this work has been dedicated to the syntheses of mixed  $C_{60}$  hexa-adducts, based on the “tether-directed” methodology developed in the first part.

Three mixed  $C_{60}$  hexa-adducts building blocks have been synthesized from *e* bis-adduct and *e,e,e* tris-adducts. The post-functionalization of the mixed  $C_{60}$  hexa-adducts building blocks has been achieved by sequential CuAAC and esterification reactions. The functionalized mixed  $C_{60}$  hexa-adducts have been obtained in very good yields. Up to three different addend zones could be predefined based on the control of the regio- and stereochemistry provided

by our methodology. The high tolerance of the CuAAC reactions to a wide variety of functions will provide an easy access to the preparation of unprecedented globular multifunctional nanomaterials with a controlled distribution of functional groups on the spherical framework.

In the last part of this work, the mixed C<sub>60</sub> hexa-adducts building blocks have been used to synthesize original fullerodendrimers. A convergent approach has been used to graft peripheral C<sub>60</sub> hexa-adducts onto a central mixed C<sub>60</sub> hexa-adduct. Our different scaffolds gave access to numerous poly- and multi-functionalized molecules. Based on our methodologies, the controlled elaboration of new nanomaterials and bioactive molecules should be easily achieved.



## Fonctionnalisation Multiple d'Hexa-adduits du C<sub>60</sub> par Chimie « Click »

### Résumé

Dans un premier temps, la bis- et tris-fonctionnalisation du C<sub>60</sub> a été effectuée par une approche macrocyclique. Différents bis- et tris-adduits du C<sub>60</sub> ont été obtenus avec de bonnes régiosélectivités. Dans un deuxième temps, la tris-fonctionnalisation du C<sub>60</sub> a été effectuée par une approche « tête directrice ». Par cette approche, un accès rapide, simple, régio- et stéréo-sélectif à des tris-adduits e,e,e approprié à la formation d'hexa-adduits d'ordre d'addition octaédrique a été obtenu. La synthèse et la séparation de tris-adduits e,e,e optiquement purs ont également été réalisées. Les dérivés polyols obtenus par déprotection des groupes silylés ouvrent de nouvelles perspectives pour la post-fonctionnalisation de multi-adduits de C<sub>60</sub>. A partir de nos méthodologies de synthèse de bis- et tris-adduits de C<sub>60</sub>, des synthons d'hexa-adduits mixtes de C<sub>60</sub> ont été préparés. La post-fonctionnalisation de ces synthons a été effectuée par réaction de cycloaddition 1,3-dipolaire de type Huisgen mais aussi par réaction d'estérification, afin d'obtenir des hexa-adduits de C<sub>60</sub> multifonctionnels. La méthodologie de synthèse d'hexa-adduits de C<sub>60</sub> multifonctionnels a été mise à profit pour l'élaboration d'édifices hyperfonctionnels comportant plusieurs hexa-adduits de C<sub>60</sub> autour d'un hexa-adduit de C<sub>60</sub> central. La post-fonctionnalisation des synthons hexa-adduits mixtes par chimie « click » donne accès à la préparation contrôlée de nouveaux nanomatériaux globulaires multifonctionnels.

**Mots clés :** [60]fullerène, chimie « click », régiochimie, stéréochimie, multi-adduits, réaction de Bingel.

### Abstract

In a first instance, the bis- and tris-functionalization of C<sub>60</sub> was performed by a macrocyclic approach. Different bis- and tris-adducts of C<sub>60</sub> were obtained with good regioselectivity. In a second instance, the tris-functionalization of C<sub>60</sub> was performed by a "tether-directed" approach. By this approach, a fast, simple, regio- and stereoselective access to C<sub>60</sub> e,e,e tris-adduct suitable for the formation of C<sub>60</sub> hexa-adducts with an octahedral addition pattern was obtained. The synthesis and the separation of optically pure e,e,e tris-adducts was also realized. The polyol derivatives obtained by deprotection of the silyl groups open new perspectives for the post-functionalization of C<sub>60</sub> multi-adducts. Using our synthesis methodologies of bis- and tris-adducts of C<sub>60</sub>, mixed hexa-adducts building blocks were prepared. The post-functionalization of these building blocks was carried out by copper catalyzed azide-alkyne 1,3-dipolar cycloadditions but also by esterification reactions to obtain multifunctional C<sub>60</sub> hexa-adducts. The methodology for the synthesis of multifunctional C<sub>60</sub> hexa-adducts was used for the elaboration of fullerodendrimers containing several C<sub>60</sub> hexa-adducts around a central C<sub>60</sub> hexa-adduct. The post-functionalization of the mixed hexa-adducts building blocks by "click" chemistry gave access to the controlled preparation of new globular and multifunctional nanomaterials.

**Keywords :** [60]fullerene, « click » chemistry, regiochemistry, stereochemistry, multi-adducts, Bingel reaction.