# Traitement et analyse d'images IRM de diffusion pour l'estimation de l'architecture locale des tissus

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### Diffusion Magnetic Resonance Imaging



Figure: 3 Tesla MRI scanner.



Figure: Brownian motion of water molecules.



Figure: Diffusion: displacement during a time t.



Figure: Free diffusion and its PDF [Einstein05].



Figure: Restricted diffusion and its PDF [Lebihan85].



### Diffusion image definition



### Diffusion image definition



We define the acquired diffusion image as:

$$E: \left| \begin{array}{c} \Omega_{\mathsf{x}} \times \Omega_{\boldsymbol{q}} \to \mathbb{R} \\ (\mathbf{x}, \mathbf{q}) \to E(\mathbf{x}, \mathbf{q}) \end{array} \right.$$

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The objective is to compute the diffusion PDF:

$$P: \left| \begin{array}{c} \Omega_{\mathsf{x}} \times \Omega_{\mathsf{p}} \to \mathbb{R} \\ (\mathsf{x}, \mathsf{p}) \to P(\mathsf{x}, \mathsf{p}) \end{array} \right.$$

Diffusion Spectrum Imaging (DSI) - Fourier transform [Wedeen00]

 $P(\mathbf{p}) = \int_{\mathbf{q}} E(\mathbf{q}) \exp(-2\pi i \mathbf{q}^T \mathbf{p}) d\mathbf{q}$ 



q-space sampling

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Figure: Human erythocytes rate for decreasing values of hematocrites [Kuchel97]. Observed in the human brain [Niendorf96].



Diffusion Spectrum Imaging (DSI) - Fourier transform [Wedeen00]

 $P(\mathbf{p}) = \int_{\mathbf{q}} E(\mathbf{q}) \exp(-2\pi i \mathbf{q}^T \mathbf{p}) d\mathbf{q}$ 



q-space sampling

#### Pro

- Complete diffusion (angular and radial profile)
- "
   No a priori on the signal
   "

#### Cons

- $\stackrel{\scriptstyle{\leftarrow}}{\sim}$  Very long acquisition time
- High gradients lead to magnetic field distortion

### Diffusion Tensor Imaging (DTI) - Gaussian assumption [Basser94]

$$E(\mathbf{q}) = \exp(-\mathbf{q}^{\mathrm{T}}\mathbf{D}\mathbf{q}) \quad \Rightarrow \quad P(\mathbf{p}) = \frac{1}{(|\mathbf{D}|(4\pi\tau)^3)^{1/2}}\exp\left(-\frac{\mathbf{p}^{\mathrm{T}}\mathbf{D}^{-1}\mathbf{p}}{4 au}
ight)$$







q-space sampling







q-space sampling















q-space sampling

-0.5<sub>0.0</sub> 0.5

0.5 ס.ס

-0.5



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ight)$$



q-space sampling

#### Pro

- $\ddot{-}$  Very short acquisition time
- Ü Well-established modality

#### Cons

- ∼ Inaccurate angular diffusion
- Simple a priori on the radial diffusion (Gaussian)

#### Several methods



#### Several methods





#### Several methods





#### Several methods





#### Several methods

Q-Ball Imaging (QBI), Diffusion Orientation Transform (DOT), etc.



q-space sampling





#### Several methods

Q-Ball Imaging (QBI), Diffusion Orientation Transform (DOT), etc.



q-space sampling

#### Pro

- $\ddot{-}$  Reduced acquisition time
- Higher angular precision than DTI

#### Cons

Simple a priori on the radial diffusion (Gaussian, Bessel, etc.)

• Various brain tissues diffuses differently, which may reveal information on cells micro-structure that composed the organic tissue.





Figure: Myelination of an axone during the childhood [www.jdaross.cwc.net]



MR radial signal

### Our aim







#### Diffusion signal estimation

• Continuous representation of the signal

#### 2) Extraction of various features of the PDF

- Algorithm
- Overview of proposed features

#### B Robust extraction of diffusion features

- Robustness to noise
- Robustness to q-space sampling

#### Diffusion signal estimation

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### Continuous representation of the signal



Figure: A sparse sampling of the *q*-space.
### Continuous representation of the signal



q-space sampling

Figure: A sparse sampling of the *q*-space.

#### Problem

Still insufficient number of samples for a Fourier transform. Which mathematical tool for the signal approximation ?

Continuous representation of the MR signal E in the following basis (Spherical Polar Fourier SPF):

$$E(\mathbf{q}) = \sum_{n=0}^{\infty} \sum_{l=0}^{\infty} \sum_{m=-l}^{l} a_{nlm} R_n(||\mathbf{q}||) y_l^m \left(\frac{\mathbf{q}}{||\mathbf{q}||}\right)$$
(1)

where  $a_{nlm}$  expansion coefficients,  $R_n$  and  $y_l^m$  are respectively are radial and angular atoms.

The basis is *orthonormal* in spherical coordinates:

$$\int_{\mathbf{q}\in\mathbb{R}^3} \left[ R_n(||\mathbf{q}||) y_l^m\left(\frac{\mathbf{q}}{||\mathbf{q}||}\right) \right] \cdot \left[ R_{n'}(||\mathbf{q}||) y_{l'}^{m'}\left(\frac{\mathbf{q}}{||\mathbf{q}||}\right) \right] d\mathbf{q} = \delta_{nn'll'mm'}$$
(2)



Figure: Real and symmetric spherical harmonics: first orders I = 0, 2, 4, 6. Blue indicates a negative value, whereas indicates red a positive value.

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Figure: Square sampled along a 5-subdivided icosahedron.

(a) 1 coefficients





Figure: Square sampled along a 5-subdivided icosahedron.

(a) 15 coefficients





Figure: Square sampled along a 5-subdivided icosahedron.

(a) 28 coefficients





Figure: Square sampled along a 5-subdivided icosahedron.

(a) 45 coefficients





Figure: Square sampled along a 5-subdivided icosahedron.

(a) 66 coefficients

Figure: Angular reconstruction along with increasing truncation order *L*.





Figure: Square sampled along a 5-subdivided icosahedron.

(a) 91 coefficients





Figure: Square sampled along a 5-subdivided icosahedron.

(a) 120 coefficients





Figure: Square sampled along a 5-subdivided icosahedron.

(a) 153 coefficients



Figure: Experimental plot [Regan06]



Figure: Experimental plot [Regan06]

Figure: Some radial atoms  $R_n$ 

### Our proposition

$$R_n\left(||\mathbf{q}||\right) = \left[\frac{2}{\gamma^{3/2}} \frac{n!}{\Gamma(n+3/2)}\right]^{1/2} \exp\left(-\frac{||\mathbf{q}||^2}{2\gamma}\right) L_n^{1/2}\left(\frac{||\mathbf{q}||^2}{\gamma}\right)$$



Figure: Radial reconstruction along with increasing truncation order N.

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Figure: Radial reconstruction along with increasing truncation order N.

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### Linear signal estimation

The representation of the signal  ${\sf E}$  as a coefficient vector  ${\sf A}$  in the basis  ${\sf M}$  is expressed as:

### $\mathbf{E} = \mathbf{M} \, \mathbf{A}$

The coefficient estimation is computed by the linear damped least square method:

$$\Rightarrow \mathbf{A} = \underset{\mathbf{A}}{\arg\min} ||\mathbf{E} - \mathbf{M}\mathbf{A}||^{2} + \lambda_{I} ||\mathbf{L}||^{2} + \lambda_{n} ||\mathbf{N}||^{2}$$
$$= (\mathbf{M}^{T}\mathbf{M} + \lambda_{I}\mathbf{L}^{T}\mathbf{L} + \lambda_{n}\mathbf{N}^{T}\mathbf{N})^{-1}\mathbf{M}^{T}\mathbf{E}$$

where M, E, A stands for:

$$\mathbf{M} = \begin{bmatrix} R_0(||\mathbf{q}_1||)y_0^0\left(\frac{\mathbf{q}_1}{||\mathbf{q}_1||}\right) & \dots & R_N(||\mathbf{q}_1||)y_L^L\left(\frac{\mathbf{q}_1}{||\mathbf{q}_1||}\right) \\ \vdots & \ddots & \vdots \\ R_0(||\mathbf{q}_{n_s}||)y_0^0\left(\frac{\mathbf{q}_{n_s}}{||\mathbf{q}_{n_s}||}\right) & \dots & R_N(||\mathbf{q}_{n_s}||)y_L^L\left(\frac{\mathbf{q}_{n_s}}{||\mathbf{q}_{n_s}||}\right) \end{bmatrix},$$
$$\mathbf{E} = (E(\mathbf{q}_1), \dots, E(\mathbf{q}_{n_s}))^T \quad \mathbf{A} = (a_{000}, \dots, a_{NLL})^T$$

### Simulation: linear least square reconstruction



Figure: Original signal



Figure: N = 1, L = 4,  $\gamma = 100$ , 1 sphere – 42 directions, PSNR: 33.337902, 30 Coefficients

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Figure: N = 3, L = 4,  $\gamma = 70$ , 3 spheres – 42 directions, PSNR: 45.172752, 45 Coefficients

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### Simulation: linear least square reconstruction



Figure: N = 5, L = 6,  $\gamma = 50$ , 10 spheres – 162 directions, PSNR: 50.255381, 168 Coefficients

### Diffusion signal estimation

Continuous representation of the signal

### 2 Extraction of various features of the PDF

- Algorithm
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### 3 Robust extraction of diffusion features

- Robustness to noise
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## Features of the PDF

Now that we have a continuous reconstruction of the diffusion signal E, how do we compute the PDF ?



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### Proposition

We are interesting in a data reduction suitable to display: *features* of the PDF.

$$\mathcal{G}(\mathsf{k}) = \int_{\mathsf{p} \in \mathbb{R}^3} \mathsf{PDF}(\mathsf{p}) \, \mathsf{H}_{\mathsf{k}}(\mathsf{p}) d\mathsf{p}$$



$$\mathcal{G}(\mathbf{k}) = \int_{\mathbf{p} \in \mathbb{R}^3} PDF(\mathbf{p}) H_{\mathbf{k}}(\mathbf{p}) d\mathbf{p}$$
(4)



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Figure: Example: ODF feature.

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Features $\mathcal{G}$	Domain ${\cal M}$
Moments	_
Return to zero	_
Anisotropy	-
Mean radial diffusion	$\mathbb{R}$
Funk-Radon Transform (FRT)	$\mathcal{S}^2$
Orientation Density Function (ODF)	$\mathcal{S}^2$
Isoradius (ISO)	$\mathcal{S}^2$
Displacement PDF	$\mathbb{R}^3$
Diffusion signal E	$\mathbb{R}^3$

Table: Overview of proposed features  $\mathcal{G}$  of the PDF.

### Moments

Moments of the diffusion PDF of order a + b + c are expressed as the scalar product between *P* and  $H^{abc}$ :

$$\mathcal{G}_{abc} = \int_{\mathbf{p} \in \mathbb{R}^3} P(\mathbf{p}) H^{abc}(\mathbf{p}) \mathrm{d}\mathbf{p} \quad \text{avec} \quad H^{abc}(\mathbf{p}) = p_x^a p_y^b p_z^c$$

These moments can be grouped as a tensor of order a + b + c, so that the DTI is a special case with a + b + c = 2.

$$\mathsf{DTI} = \begin{pmatrix} \mathcal{G}_{200} & \mathcal{G}_{110} & \mathcal{G}_{101} \\ \mathcal{G}_{110} & \mathcal{G}_{020} & \mathcal{G}_{011} \\ \mathcal{G}_{101} & \mathcal{G}_{011} & \mathcal{G}_{002} \end{pmatrix}$$



The Funk-Radon transform approximates the ODF. It is used by HARDI methods such as the Q-Ball Imaging.

The FRT feature is written as:

$$\mathcal{G}(\mathbf{k}) = \int_{\mathbf{p} \in \mathbb{R}^3} H_{\mathbf{k}}(\mathbf{p}) P(\mathbf{p}) \mathrm{d}\mathbf{p}$$

where  $H_{\mathbf{k}}$  is the associated projection function at point  $\mathbf{k} \in \mathcal{S}^2$ . Let  $q' \in \mathbb{R}$  be the sampling radius of the q-sphere and  $\mathbf{p} = p\mathbf{r}$ , so that [Tuch04]:

$$H_{\mathbf{k}}(\mathbf{p}) = 2\pi q' J_0(2\pi q' p) \delta \left(1 - \mathbf{r} \cdot \mathbf{k}\right)$$

# Funk-Radon Transform (FRT)



Figure: Reproduction of the QBI method by our approach.

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The diffusion orientation density function (ODF) is the projection of the PDF on the unit sphere. This feature is expressed as:

$$\mathcal{G}(\mathsf{k}) = \int_{\mathbf{p} \in \mathbb{R}^3} P(\mathbf{p}) \mathcal{H}_{\mathsf{k}}(\mathbf{p}) \mathrm{d}\mathbf{p}$$

where  $H_{\mathbf{k}}$  is

$$H_{\mathbf{k}}(\mathbf{p}) = \delta(1 - \mathbf{r} \cdot \mathbf{k})$$



Figure: ODF approximation at low, medium and high resolution.



(a) DTI (Basser94)

Figure: Brain white matter: ODF overlayed on a GFA map. DTI(a) and QBI(b) were computed with  $b = 3000 \text{ s/mm}^2$ . Our method (c) shows the obtained ODF with b = 1000 and  $3000 \text{ s/mm}^2$ .

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The anisotropy is a scalar measure which is useful to have hindsight on the wiring structure of the nerve fibers across the brain.

The generalized fractional anisotropy feature, which generalizes the fractional anisotropy (FA), is expressed as:

$$\mathsf{GFA}(\mathcal{G}) = \frac{\mathsf{std}(\mathcal{G})}{\mathsf{rms}(\mathcal{G})} = \sqrt{\frac{\int_{\mathbf{k}\in\mathcal{S}^2} (\mathcal{G}(\mathbf{k}) - \langle \mathcal{G} \rangle)^2 \mathrm{d}\mathbf{k}}{\int_{\mathbf{k}\in\mathcal{S}^2} \mathcal{G}(\mathbf{k})^2 \mathrm{d}\mathbf{k}}}$$

where  $\mathcal{G} : \mathcal{S}^2 \to \mathbb{R}$  is a spherical feature of the PDF (*e.g.*  $\mathcal{G} = \mathsf{FRT,ODF,ISO}$ , *etc.*).



Figure: Anisotropy of several spherical features. (a)-(c) Anisotropy based on the DTI, QBI et true ODF.

Features $\mathcal G$	Domain ${\cal M}$
Moments	_
Return to zero	_
Anisotropy	_
Mean radial diffusion	$\mathbb{R}$
Funk-Radon Transform (FRT)	$\mathcal{S}^2$
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#### Problem

Acquisition noise not Gaussian but Rician.

Nontheless most methods use least square estimation !

 $\Rightarrow$  Bias especially at low SNR values (high *b* values).



### Variational framework

Robustly estimate and regularize the SPF coefficients by minimizing the functional energy:

$$\min_{A} \left\{ \int_{\Omega_{E}} \left[ \sum_{k}^{n_{s}} \psi(\hat{\mathbf{E}}_{k}) \right] + \alpha_{r} \varphi(||\nabla \mathbf{A}||) d\Omega_{E} \right\}, \text{ with } \hat{\mathbf{E}} = \mathbf{M} \mathbf{A}$$
(5)

The best fitting coefficients A are computed with a gradient descent coming from the Euler-Lagrange derivation of the energy. This leads to a set of *multi-valued* partial derivate equation.

$$\begin{cases} \mathbf{A}_{t=0} = U_0 \\ \frac{\partial \mathbf{A}_j}{\partial t} = \sum_k^{n_s} \mathbf{M}_{k,j} \psi'(\hat{\mathbf{E}}_k) + \alpha_r \operatorname{div}(\varphi(||\nabla \mathbf{A}||)) \end{cases}$$
(6)  

$$\underbrace{\mathbf{A}_{j}}_{\mathbf{A}_j} = \sum_k^{n_s} \mathbf{M}_{k,j} \psi'(\hat{\mathbf{E}}_k) + \alpha_r \operatorname{div}(\varphi(||\nabla \mathbf{A}||))$$
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(9)  

$$\underbrace{\mathbf{A}_{j}}_{\mathbf{A}_j} = \sum_{k=1}^{n_s} \mathbf{A}_{j} + \sum_{k=1}^{n_s} \mathbf$$

### • $\psi$ -likelihood function adapted to MRI noise law:

The best  $\psi$  function is the one specific to MR scanners, ie. Rice distribution:

$$p(E|\hat{E},\sigma) = \frac{E}{\sigma^2} \exp\left(\frac{-(E^2 + \hat{E}^2)}{2\sigma^2}\right) I_0\left(\frac{E \cdot \hat{E}}{\sigma^2}\right)$$
(7)

We seek  $\hat{E}$  which maximizes a posteriori (MAP) the log-posterior probability [Basu06]

$$\log p(\hat{E}|E) = \log p(E|\hat{E}) + \log p(\hat{E}) - \log p(E)$$
(8)

Consequently the pointwise likelihood is

1

$$\log p(E|\hat{E},\sigma) = \log \frac{E}{\sigma^2} - \frac{(E^2 + \hat{E}^2)}{2\sigma^2} + \log I_0\left(\frac{E \cdot \hat{E}}{\sigma^2}\right) = \psi(\hat{E})$$
(9)

### Advantages: regularity

• Ensure a global regularity of the SPF field:  $\varphi(||\nabla \mathbf{A}||) = \varphi(\sum_{nlm} ||\nabla A_{nlm}||)$ 

 $\varphi$  is a contour preserving function, widely used in image processing. Example of possible regularization function  $\varphi$ 



# Simulation: validation on synthetical data (likelihood)



Figure: Synthetic phantom of networks of crossing fibers. Performances of likelihood functions on increasing levels of noise.

### Simulation: Rician vs Gaussian likelihood function



### Simulation: energy minimization



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Figure: Comparison of GFA on region of corpus callosum and lateral ventricles.



# Fiber-tracking

GFA



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GFA



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GFA



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### Fiber-tracking

PDE

#### GFA



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### Questions





### Questions

• Which sampling distribution gives the best results ?



### Questions

- Which sampling distribution gives the best results ?
- Is it possible to unify the sparse sampling in the literature in one model ? [Assaf05, Özarslan06, Wu07, Khachaturian07, Assemlal-et.al08, Assemlal-et.al09]

$$f_{\mathsf{x}}(\eta) = \frac{q_{\mathsf{x}}^{\eta}}{\sum_{i=1}^{n_b} q_i^{\eta}} n_{\mathsf{s}}, \quad \text{and} \quad q_i(\beta) = \left(\frac{i-1}{n_b-1}\right)^{\beta} (q_{\max} - q_{\min}) + q_{\min}$$

$$f_{\scriptscriptstyle X}(\eta) = rac{q_{\scriptscriptstyle X}^{\eta}}{\sum_{i=1}^{n_b} q_i^{\eta}} n_{\scriptscriptstyle S}, \hspace{1em} ext{and} \hspace{1em} q_i(eta) = \left(rac{i-1}{n_b-1}
ight)^{eta} (q_{max}-q_{min}) + q_{min}$$



Figure: Condition number  $C = ||\mathbf{M}_{reg}||_{\infty} ||\mathbf{M}_{reg}^{-1}||_{\infty}$ . The lower C is, the more stable the reconstruction is. Data simulates crossing fibers diffusion signal.  $L = 4, N = 3, n_b \ge 3$ . (d)  $\lambda_n = 10^{-4}, \lambda_l = 10^{-6}$ .



(a)  $n_s = 300$ No regularization.

Figure: Condition number  $C = ||\mathbf{M}_{reg}||_{\infty} ||\mathbf{M}_{reg}^{-1}||_{\infty}$ . The lower C is, the more stable the reconstruction is. Data simulates crossing fibers diffusion signal.  $L = 4, N = 3, n_b \ge 3$ . (d)  $\lambda_n = 10^{-4}, \lambda_l = 10^{-6}$ .



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HE. Assemlal (GREYC)

Diffusion MRI

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#### Diffusion signal estimation

• Continuous representation of the signal

#### 2) Extraction of various features of the PDF

- Algorithm
- Overview of proposed features

#### B Robust extraction of diffusion features

- Robustness to noise
- Robustness to q-space sampling

## Conclusion: 1-MR signal approximation



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Diffusion MRI

Features $\mathcal{G}$	Domain ${\cal M}$
Moments	_
Return to zero	-
Anisotropy	-
Mean radial diffusion	$\mathbb{R}$
Funk-Radon Transform (FRT)	$\mathcal{S}^2$
Orientation Density Function (ODF)	$\mathcal{S}^2$
Isoradius (ISO)	$\mathcal{S}^2$
Displacement PDF	$\mathbb{R}^3$
Diffusion signal E	$\mathbb{R}^3$

Table: Overview of proposed features  $\mathcal{G}$  of the PDF.

## Conclusion: 3-Robustness

### Robustness to noise



### Robustness to sampling distribution

	$\eta = -2$	$\eta = -1$	$\eta = 0$	$\eta = 1$	$\eta = 2$	$\eta = 3$
$n_b = 2$	•	•	•			-
$n_b = 5$						



#### Thank you for your attention.

# Radial eigenfunctions

### Proposition

We propose the use of Gauss-Laguerre functions:

$$R_n(||\mathbf{q}||) = \left[\frac{2}{\gamma^{3/2}} \frac{n!}{\Gamma(n+3/2)}\right]^{1/2} \exp\left(-\frac{||\mathbf{q}||^2}{2\gamma}\right) L_n^{1/2}\left(\frac{||\mathbf{q}||^2}{\gamma}\right), \quad (10)$$

where  $\gamma$  is the scale factor.

Let  $L_n^k$  be a generalized Laguerre polynomial:

$$L_{n}^{k} = \frac{1}{n!} \sum_{i=0}^{n} \frac{n!}{i!} \binom{k+n}{n-i} (-x)^{i}$$
(11)

The Gauss attenuation comes from the normalization of these polynomials:

$$\int_{0}^{\infty} e^{-k} x^{k} L_{n}^{k}(x) L_{n'}^{k}(x) dx = \frac{(n+k)!}{n!} \delta_{nn'}$$
(12)



Figure: Power spectrum along with increasing radial truncation order.

## Runge-Kutta



Figure: Comparaison de méthodes de suivi de fibres C, définie par  $y' = \sin(t)^2 * y$ .

## PDE initialization



Figure: Influence de l'initialisation  $A_0$  sur la convergence de la méthode de descente du gradient. Les données sont bruités telles que PSNR(Originale,Bruité)=19.5 dB.

Nom de la fonction	$s  ightarrow arphi(s),  s \in \mathbb{R}$
Tikhonov	<i>s</i> <sup>2</sup>
Perona-Malik	$1-\exp(-s^2/\kappa^2)$
Surfaces minimales	$2\sqrt{1+s^2}-2$
German-McClure	$s^2/(1+s^2)$
Variation totale	S
Green	$2\log(\cosh(s))$

Table: Liste de quelques fonctions de régularisation  $\varphi$  possibles. Tableau adapté de [Tschumperlé02].



Figure: Histogramme du bruit dans les images d'IRM de diffusion.

Domaine ${\cal M}$	$\mathbb{R}$	$\mathbb{R}^3$	$S^2$
Tps d'exécution	9 s	$\sim$ 360 h	22 s

Table: Temps d'exécution de la construction d'une caractéristique dans l'ensemble de son domaine  $\{h_{nlm}^{\mathbf{k}}, \forall \mathbf{k} \in \mathcal{M}\}$ .

Nombre de valeurs de $\zeta$	1	16	224
Tps d'exécution	12 s	21 s	5 min

 Table: Temps d'exécution pour l'estimation du signal de diffusion dans la base

 SPF.

Domaine ${\cal M}$	$\mathbb{R}$	$\mathbb{R}^3$	$\boldsymbol{\mathcal{S}}^2$
Tps d'exécution	5 s	21 s	9 s

Table: Temps d'exécution pour l'extraction d'une caractéristique.

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### Return to zero probability

Scalar feature of free / restriction diffusion.

$$\mathcal{G} = P(0) = \int_{\mathbf{p} \in \mathbb{R}^3} P(\mathbf{p}) H(\mathbf{p}) d\mathbf{p} \quad \text{avec} \quad H(\mathbf{p}) = \delta(\mathbf{p})$$



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Figure: Free diffusion (a-b) and restricted (c-d) in a voxel.

## Average radial diffusion

Radial diffusion robust to anisotropy.



### Feature which reproduces the results of the DOT method [Özarslan06].

