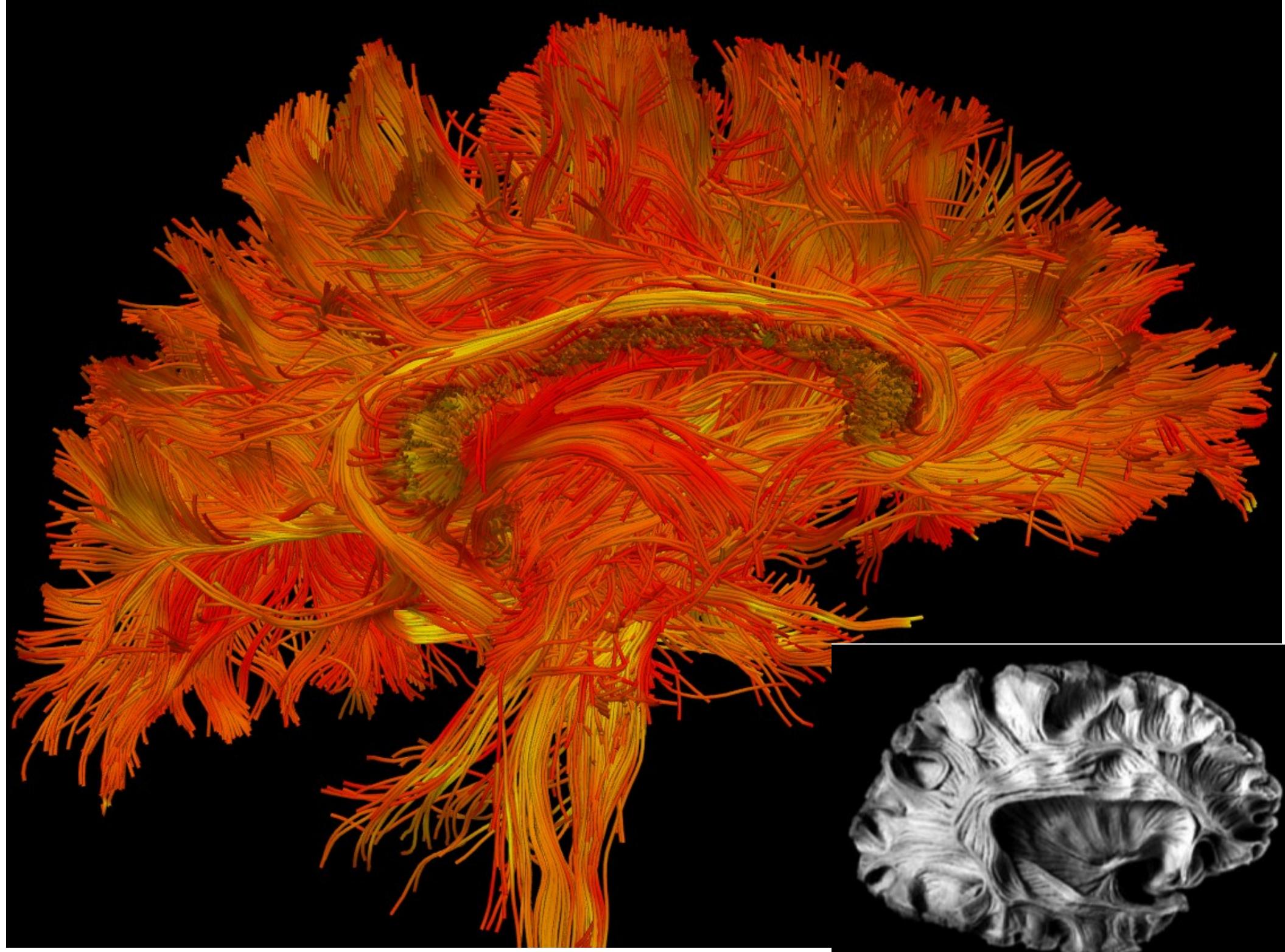


# Neural Tractography Using an Unscented Kalman Filter

James Malcolm

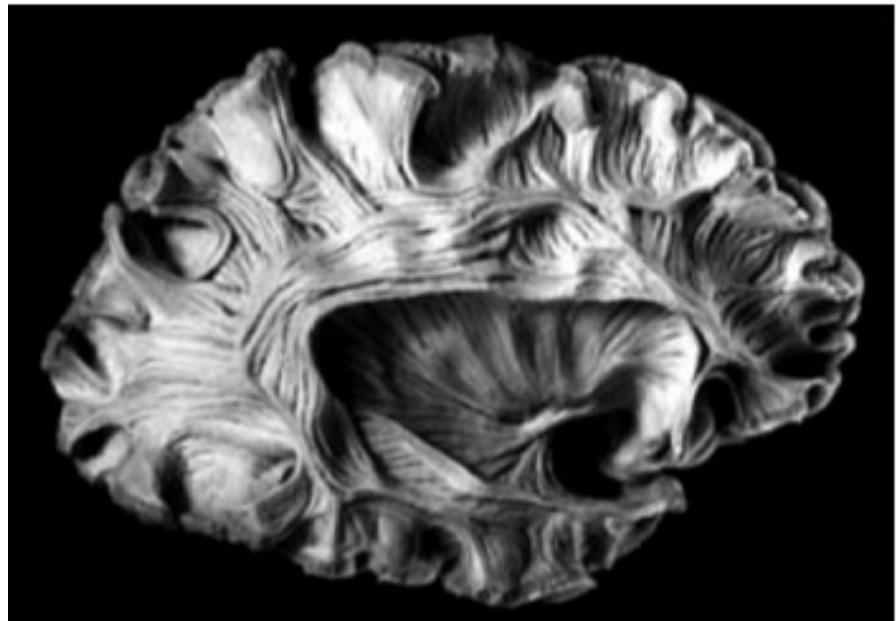
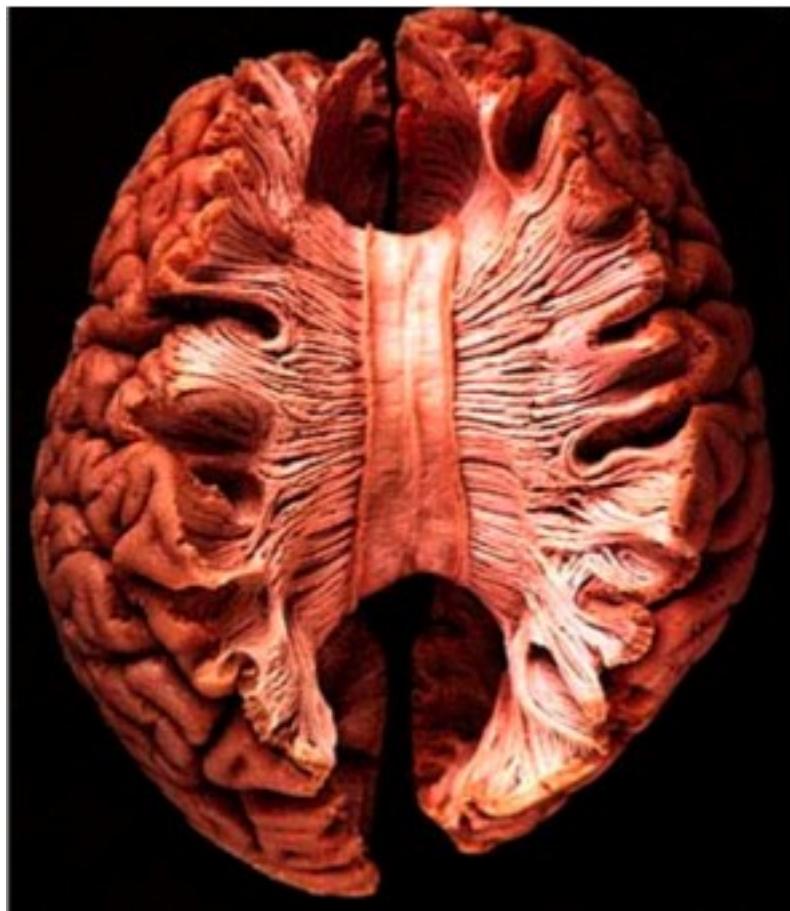
PhD Defense  
21 Oct 2010



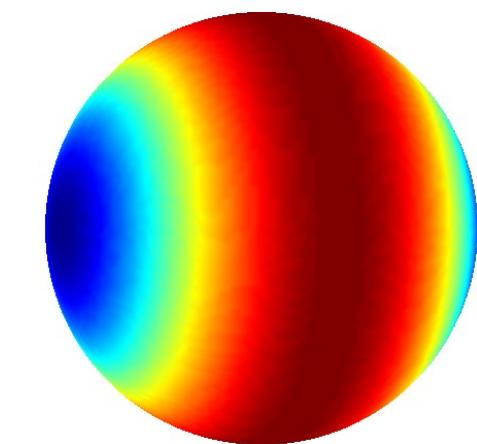
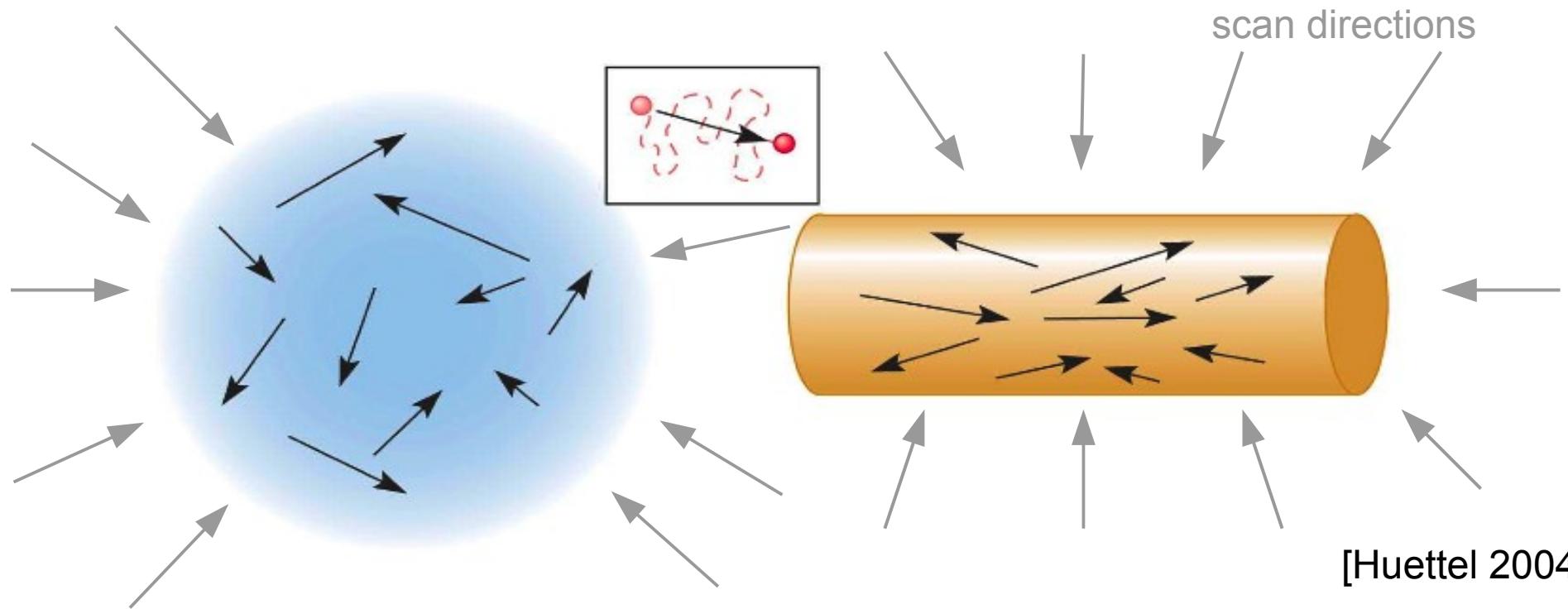
# background

J. G. Malcolm, Y. Rathi, and C.-F. Westin. “Processing and visualization of diffusion MRI.” In T. Deserno, editor, *Recent Advances in Biomedical Image Processing and Analysis*, chapter 16, pages 387–410. Springer, 2011.

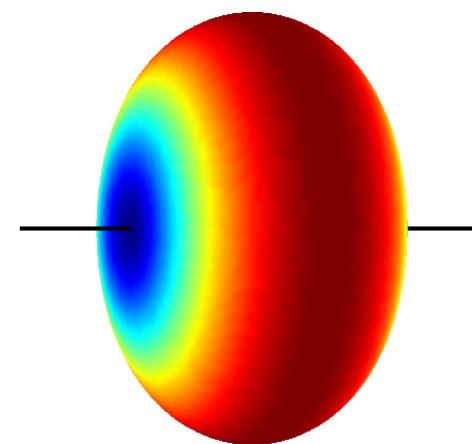
# neural fibers



[Williams<sup>4</sup> 97]



isotropic scanner signal

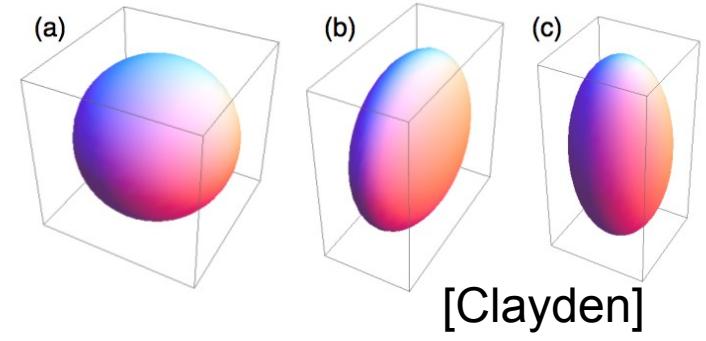


directed signal

# imaging techniques

- Diffusion tensor imaging (DTI)  
single-tensor

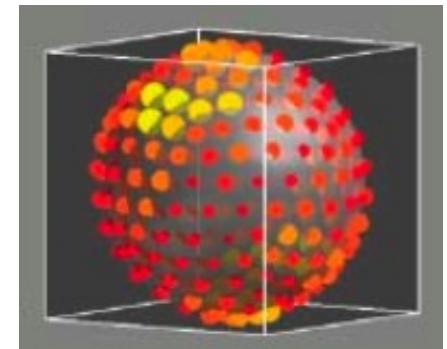
[Basser 94]



[Clayden]

- High Angular Resolution (QBI)  
multi-fiber

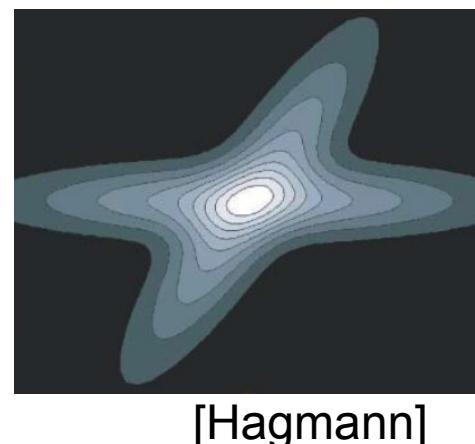
[Tuch 02]



[Tuch]

- Diffusion Spectrum (DSI)  
radial information

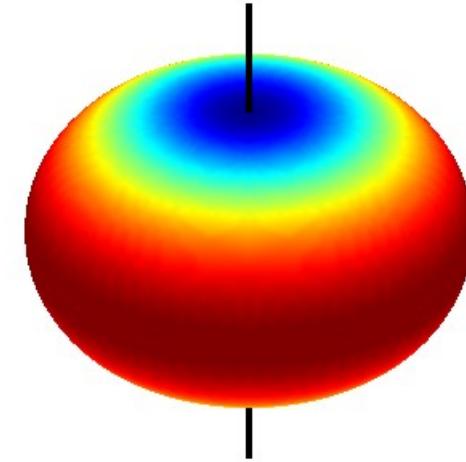
[Wedgeen 05, Hagmann 05]



[Hagmann]



one neural fiber  
0.2 to 20 microns

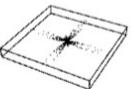
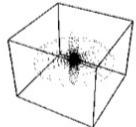
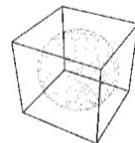


one scanned voxel  
 $1.7 \times 1.7 \times 1.7$  mm

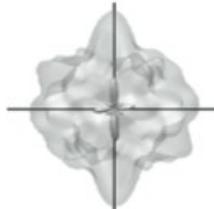
# voxel reconstruction



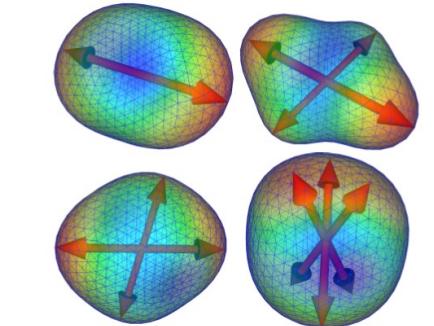
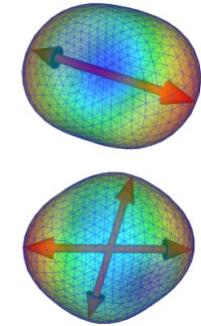
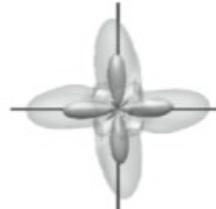
tensor mixtures [Tuch 02]



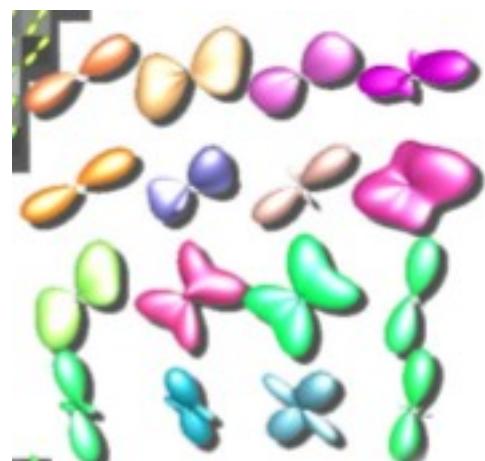
PAS [Jansons 03], MESD [Alexander 05]



SD [Tournier 04]



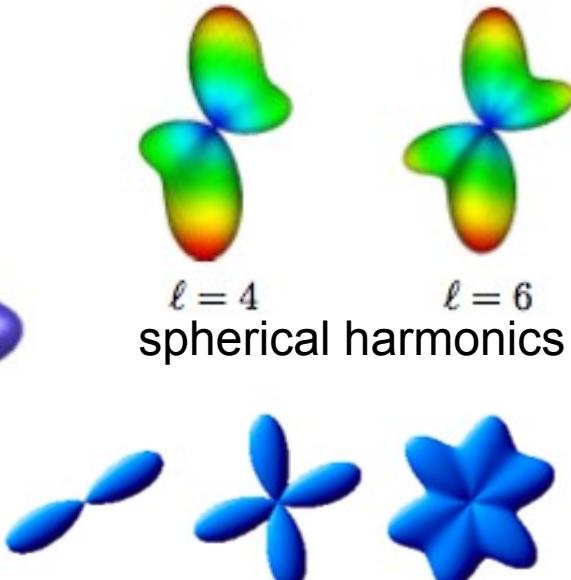
HOT [Hlawitschka 05]



VMF [Kumar 08]



DOT [Ozarslan 05]



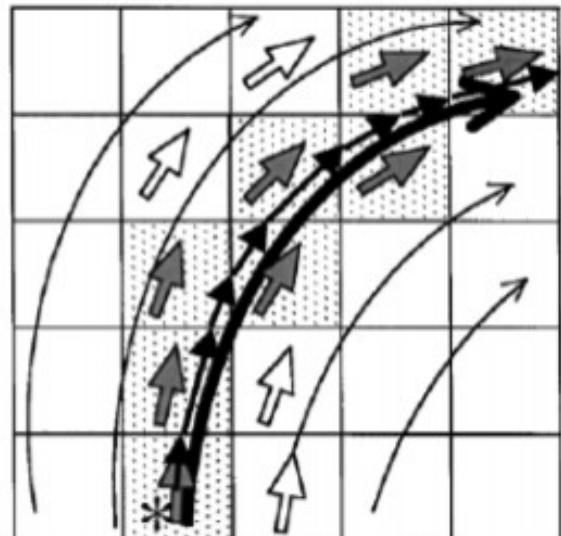
$\ell = 4$        $\ell = 6$        $\ell = 8$   
spherical harmonics [Descoteaux 07]



Wishart [Jian 07]

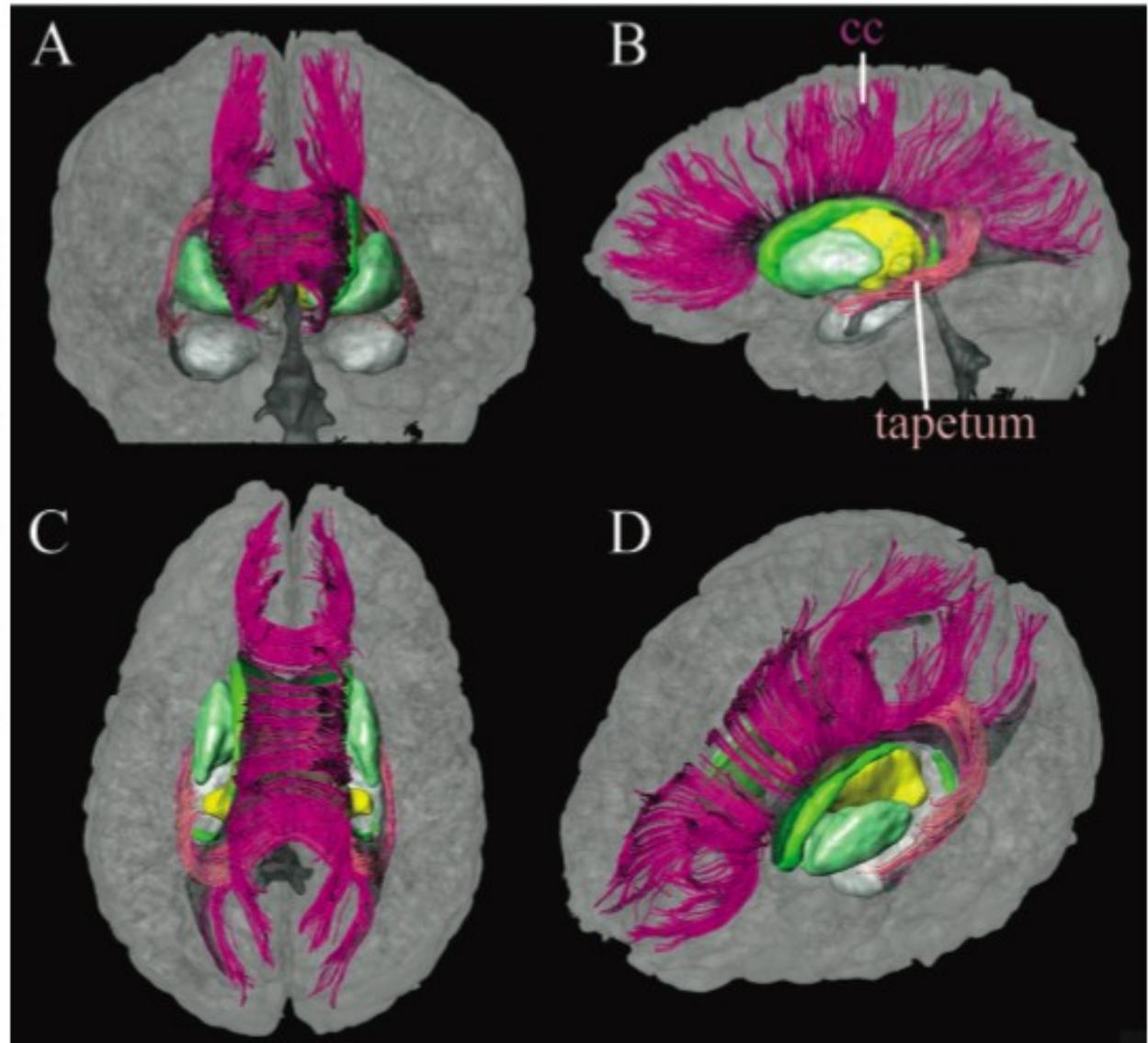
# deterministic tractography

streamlin  
e



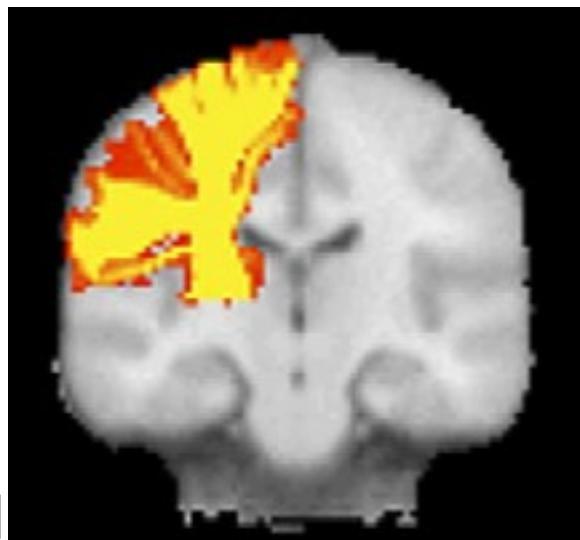
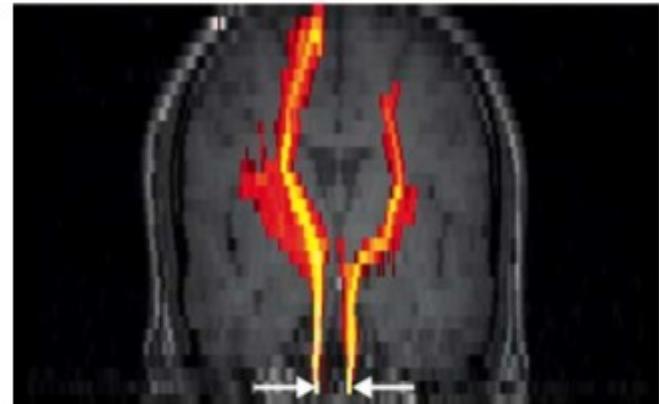
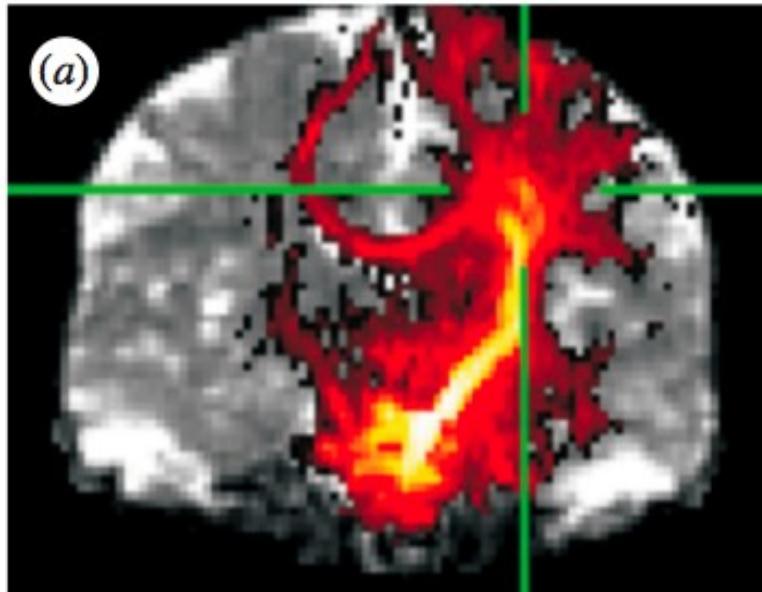
[Mori 99]

[Conturo 99, Basser 00]



[Wakana 03]

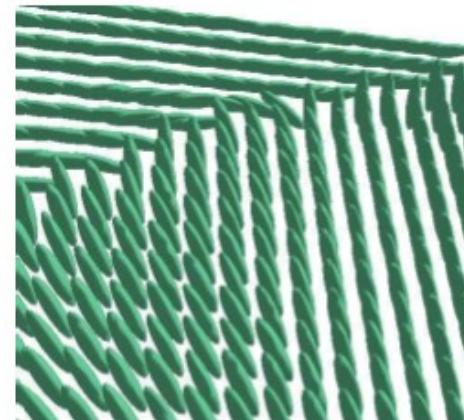
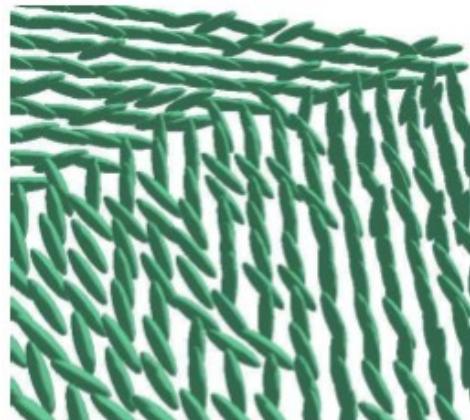
# probabilistic tractography



[Friman 06]

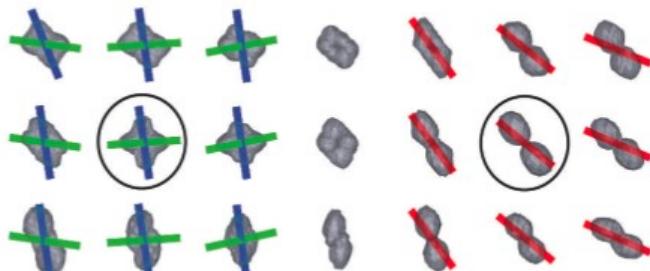
[Behrens 07]

# spatial regularization

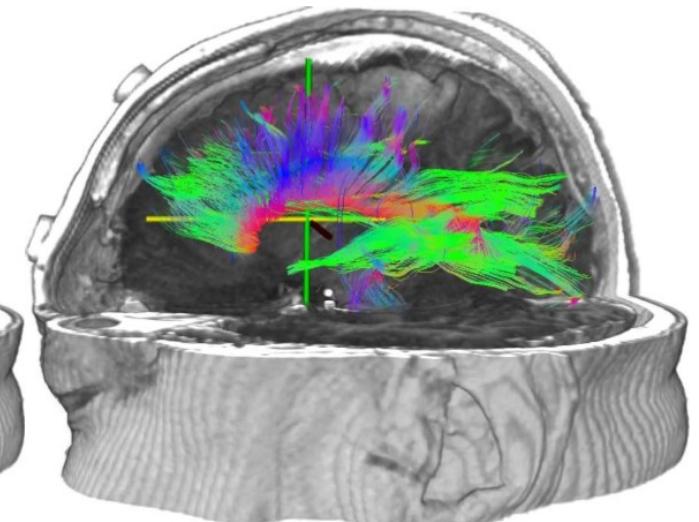
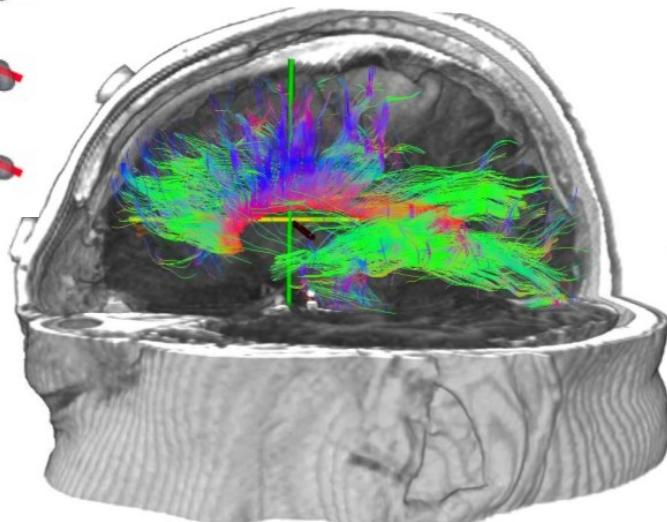


pre/post-processing

[Tschumperle 03]



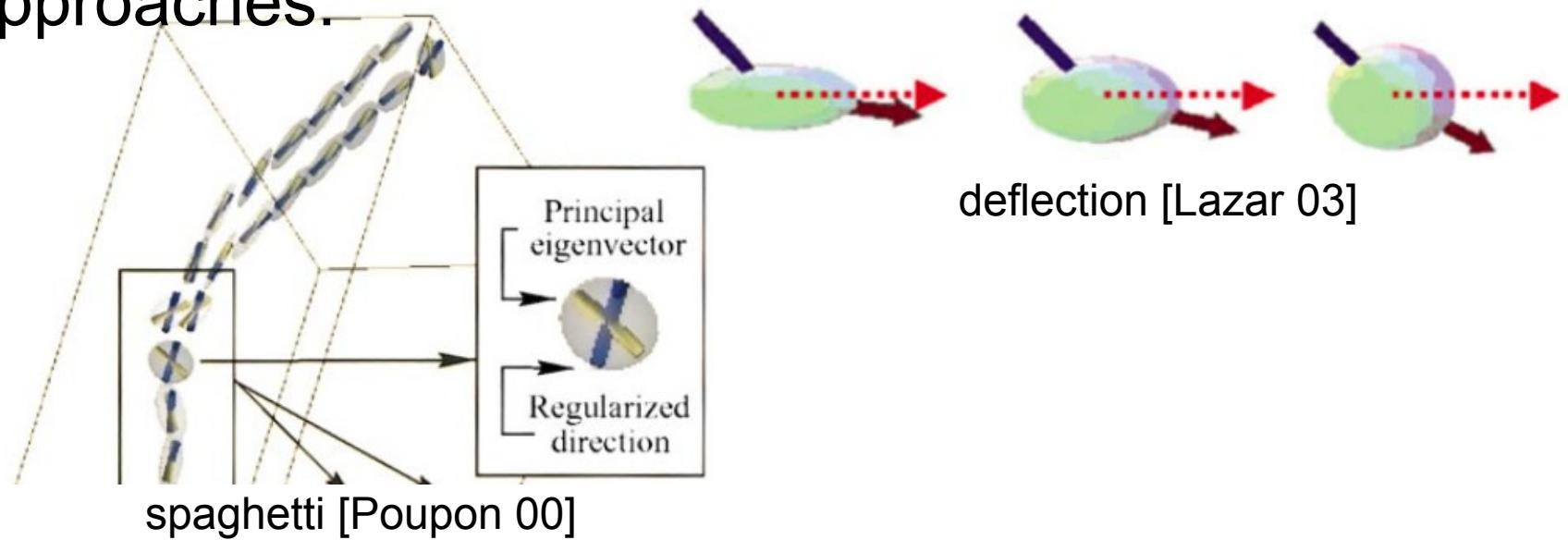
[Savadjiev 07]



[Fillard 07]

# tract regularization

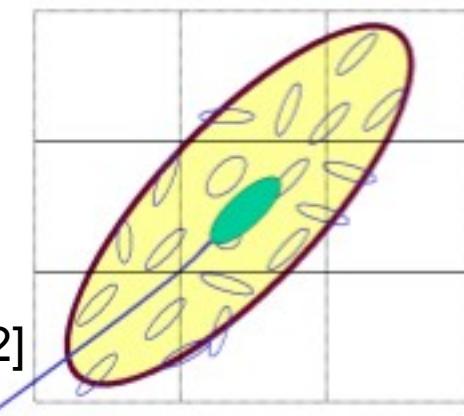
deterministic  
approaches:



spaghetti [Poupon 00]

deflection [Lazar 03]

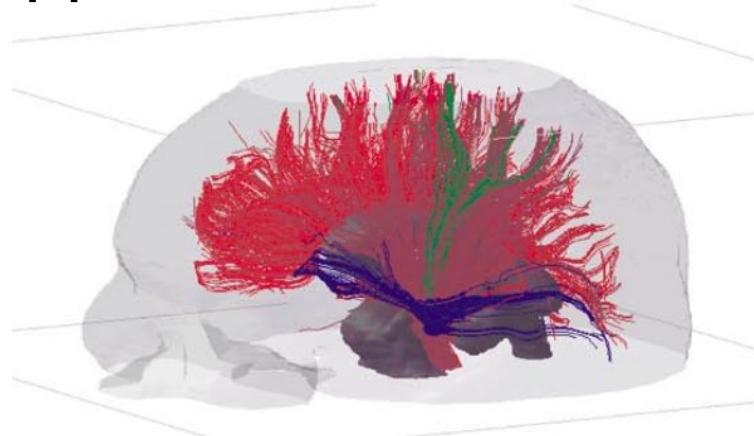
[Zhukov 02]



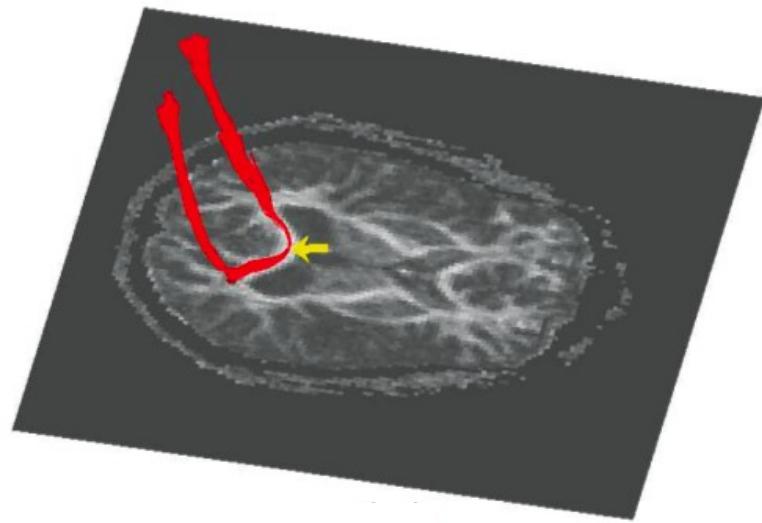
single-tensor  
path  
regularization

# tract regularization

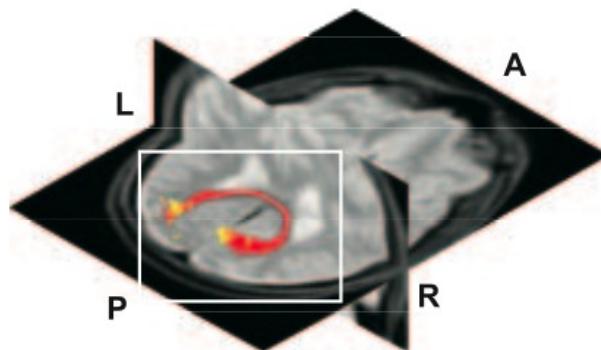
probabilistic  
approaches:



linear Kalman filter [Gossl 02]



particle filtering [Zhang 07]



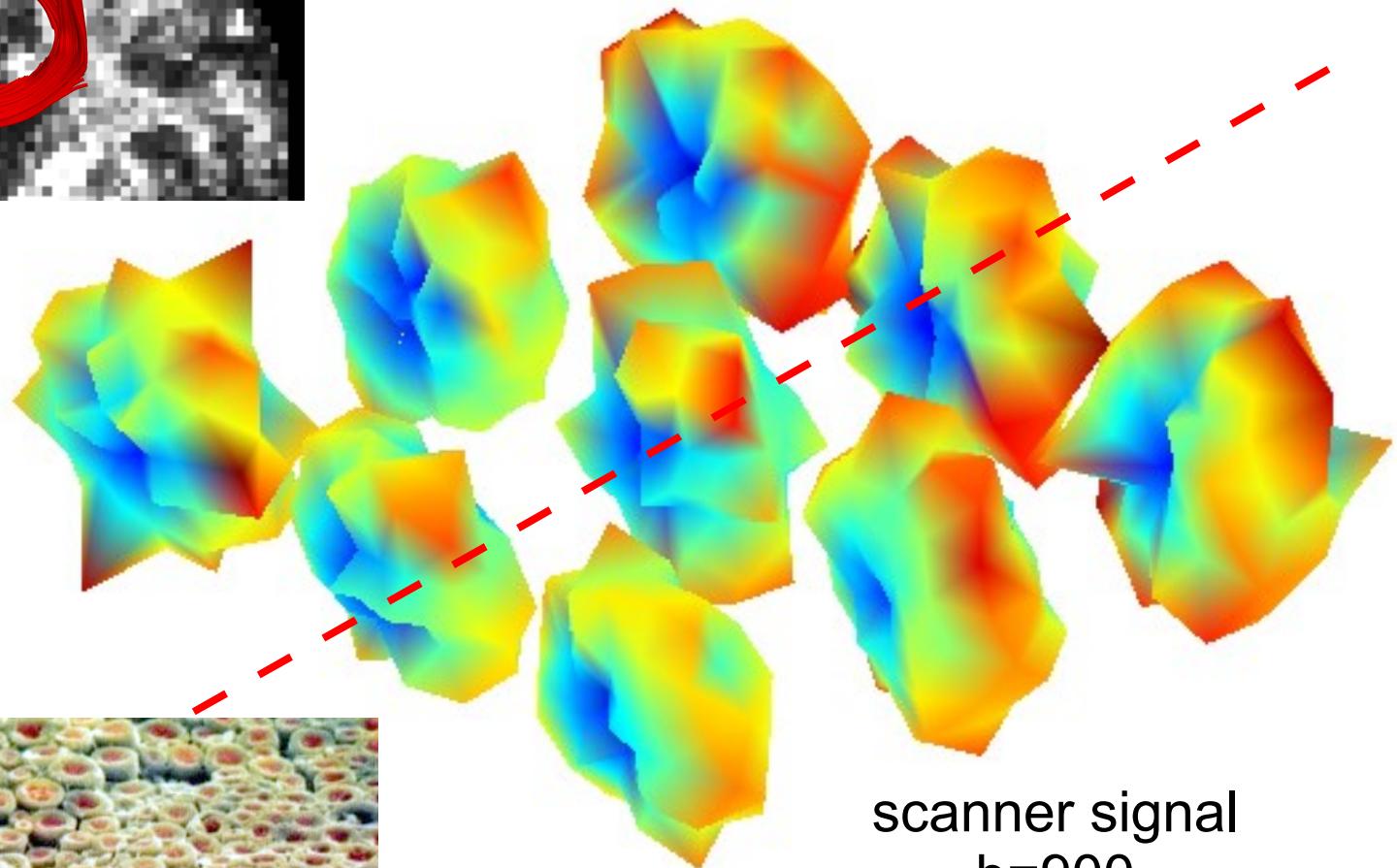
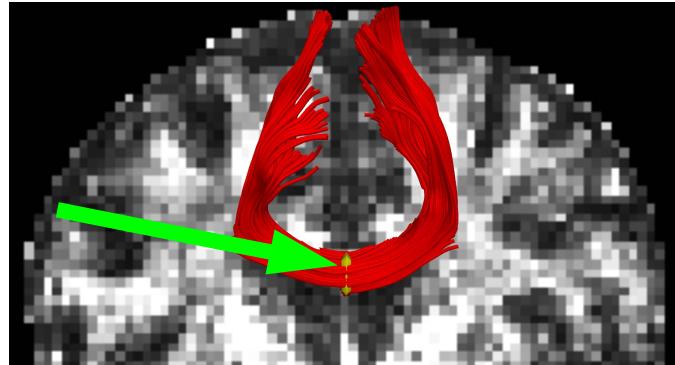
stochastic [Bjornemo 02]

single-tensor  
path  
regularization

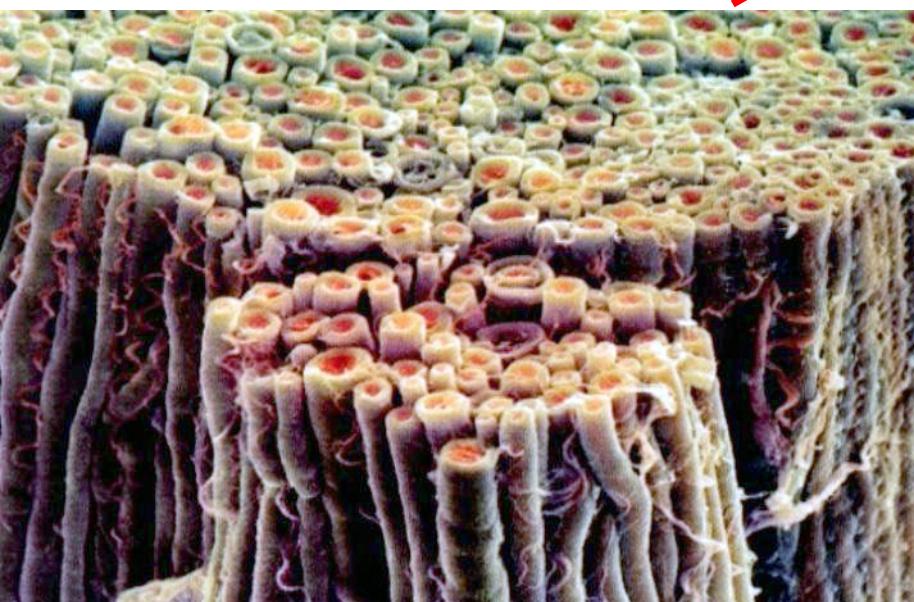
# method: the model

Y. Rathi, J. G. Malcolm, O. Michailovich, C.-F. Westin, M. E. Shenton, and S. Bouix. Tensor-kernels for simultaneous fiber model estimation and tractography. *Magnetic Resonance in Medicine*, 64(1):138–148, 2010.

J. G. Malcolm, M. E. Shenton, and Y. Rathi. Neural tractography using an unscented Kalman filter. In *Information Processing in Medical Imaging (IPMI)*, pages 126–138, 2009.



scanner signal  
 $b=900$   
51 directions  
1.7mm isotropic voxel  
17 minute scan



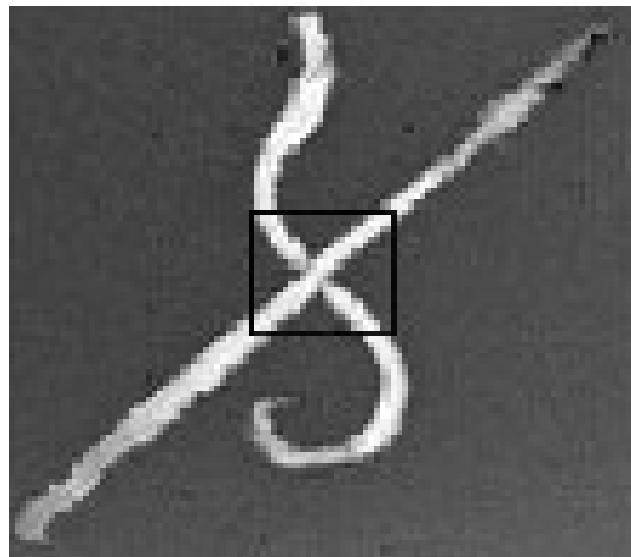
[Bartzokis]

# DTI

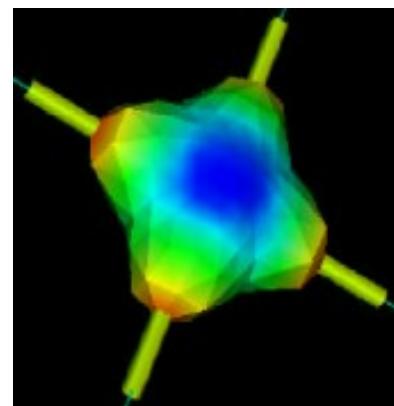
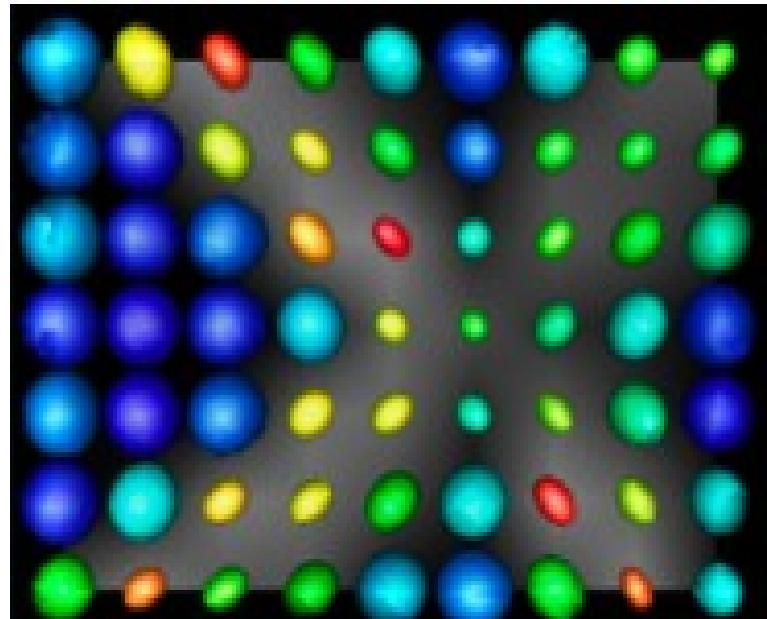
$$S(\mathbf{u}) = s_0 e^{-b \mathbf{u}^T D \mathbf{u}}$$

$D$	diffusion tensor	6 parameters
$\mathbf{u}$	unit direction	
$b$	acquisition constant	
$s_0$	null signal ( $b=0$ )	

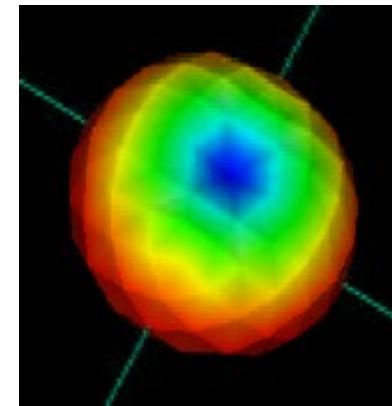
# single-tensor limitations



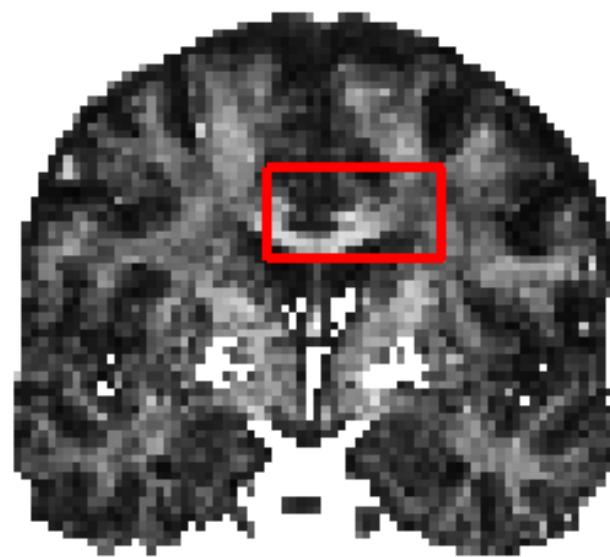
rat spinal nerves

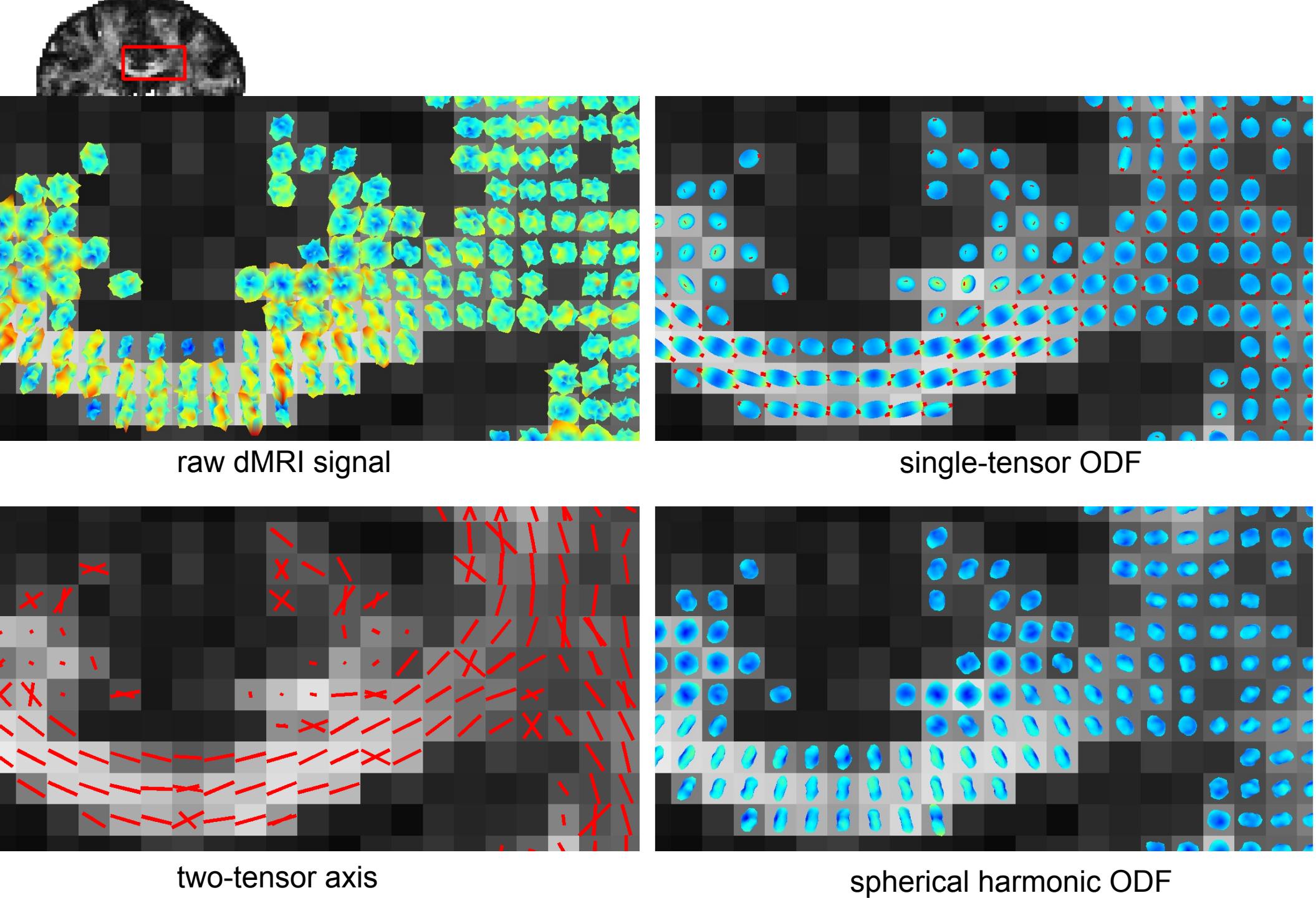


true ODF



single-tensor ODF





# multi-tensor signal model

$$S(\mathbf{u}) = s_0 \sum_j w_j e^{-b \mathbf{u}^T D_j \mathbf{u}}$$

$D_j$  diffusion tensor

$\mathbf{u}$  unit direction

$w_j$  convex weights

$b$  acquisition constant

$s_0$  null signal ( $b=0$ )

# model assumptions

*...for now*

Two fibers

Fixed volume fractions

Tensors are elliptic or isotropic

# model parameters

for two fibers...

...two principal directions     $\mathbf{m} \in \mathbb{R}^3$

...two primary eigenvalues     $\lambda_1 \in \mathbb{R}$

...two minor eigenvalues     $\lambda_2 \in \mathbb{R}$

5 + 5 = 10 parameters

# model parameters

for two fibers...

...two principal directions     $\mathbf{m} \in \mathbb{R}^3$

...two primary eigenvalues     $\lambda_1 \in \mathbb{R}$

...two minor eigenvalues     $\lambda_2 \in \mathbb{R}$

**5 + 5 = 10 parameters**

$$S(\mathbf{u}) = 0.5 s_0 e^{-b \mathbf{u}^T D_1 \mathbf{u}} + 0.5 s_0 e^{-b \mathbf{u}^T D_2 \mathbf{u}}$$

$$D_1 = \lambda_{11} \mathbf{m}_1 \mathbf{m}_1^T + \lambda_{21} (\mathbf{p} \mathbf{p}^T + \mathbf{q} \mathbf{q}^T)$$

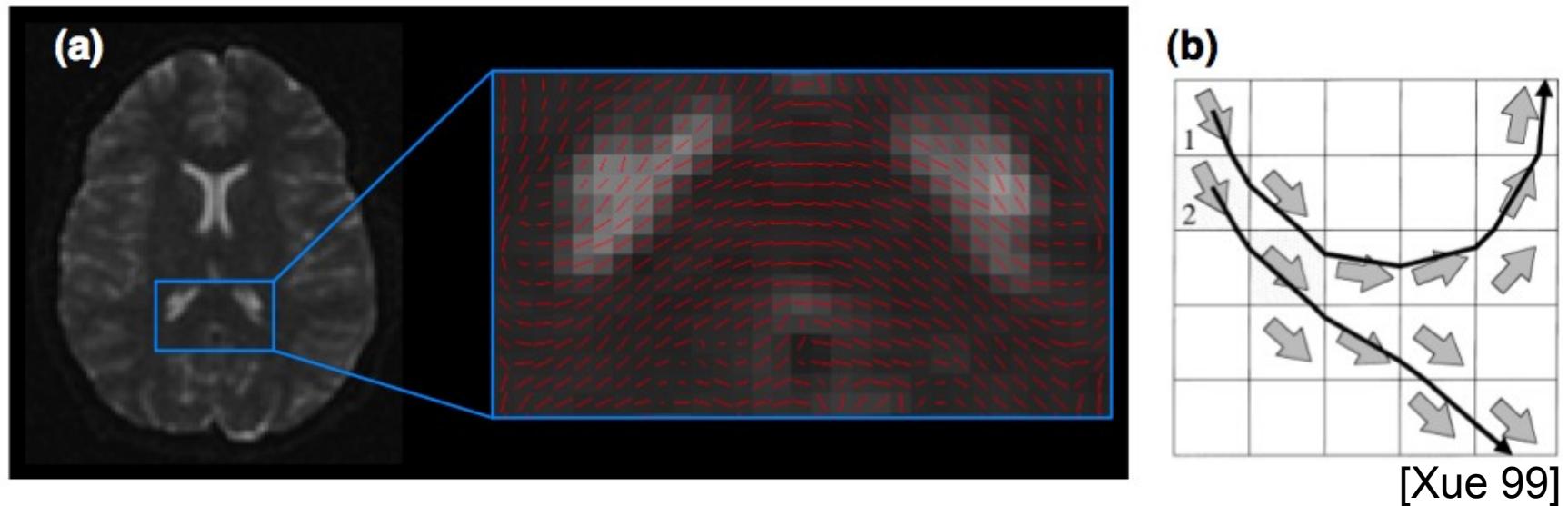
eigenvectors:  $\mathbf{m}, \mathbf{p}, \mathbf{q}$

# method: estimating the model

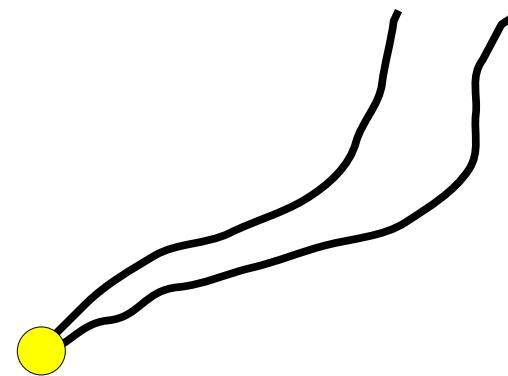
J. G. Malcolm, O. Michailovich, S. Bouix, C.-F. Westin, M. E. Shenton, and Y. Rathi. A filtered approach to neural tractography using the Watson directional function. *Medical Image Analysis*, 14:58–69, 2010.

J. G. Malcolm, M. E. Shenton, and Y. Rathi. Filtered multi-tensor tractography. *IEEE Trans. on Medical Imaging*, 29:1664–1675, 2010.

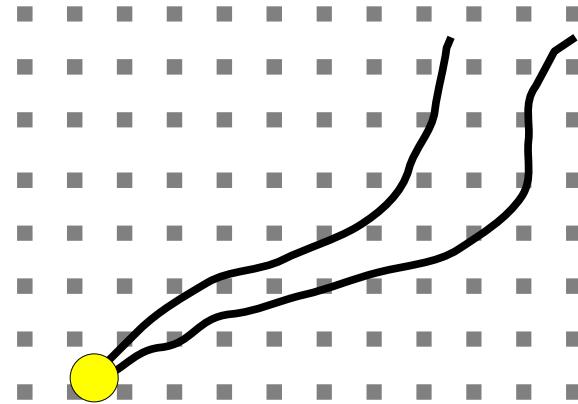
# independent estimation



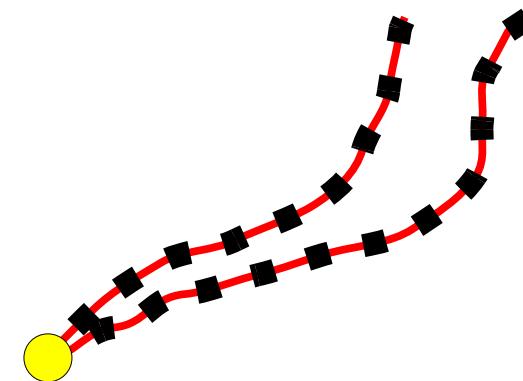
# the system: a fiber



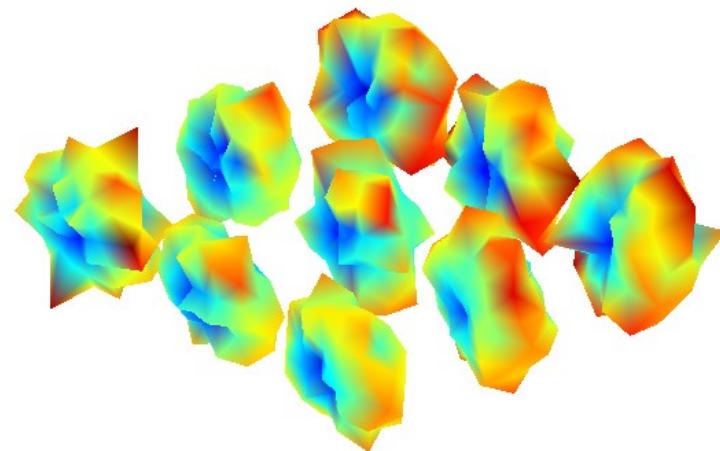
**independent  
process**



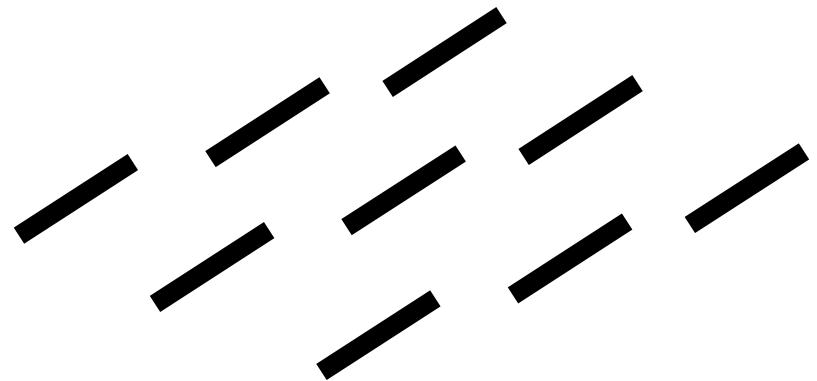
**causal  
process**



# model-based filtering



scanner  
measurement



underlying  
model

objectives:

- estimate model from measurements
- suppress noise

# notation

$\boldsymbol{x}_t$  state of system at time  $t$   
state = “model parameters”

$\boldsymbol{y}_t$  what you see at time  $t$   
observation, measurement

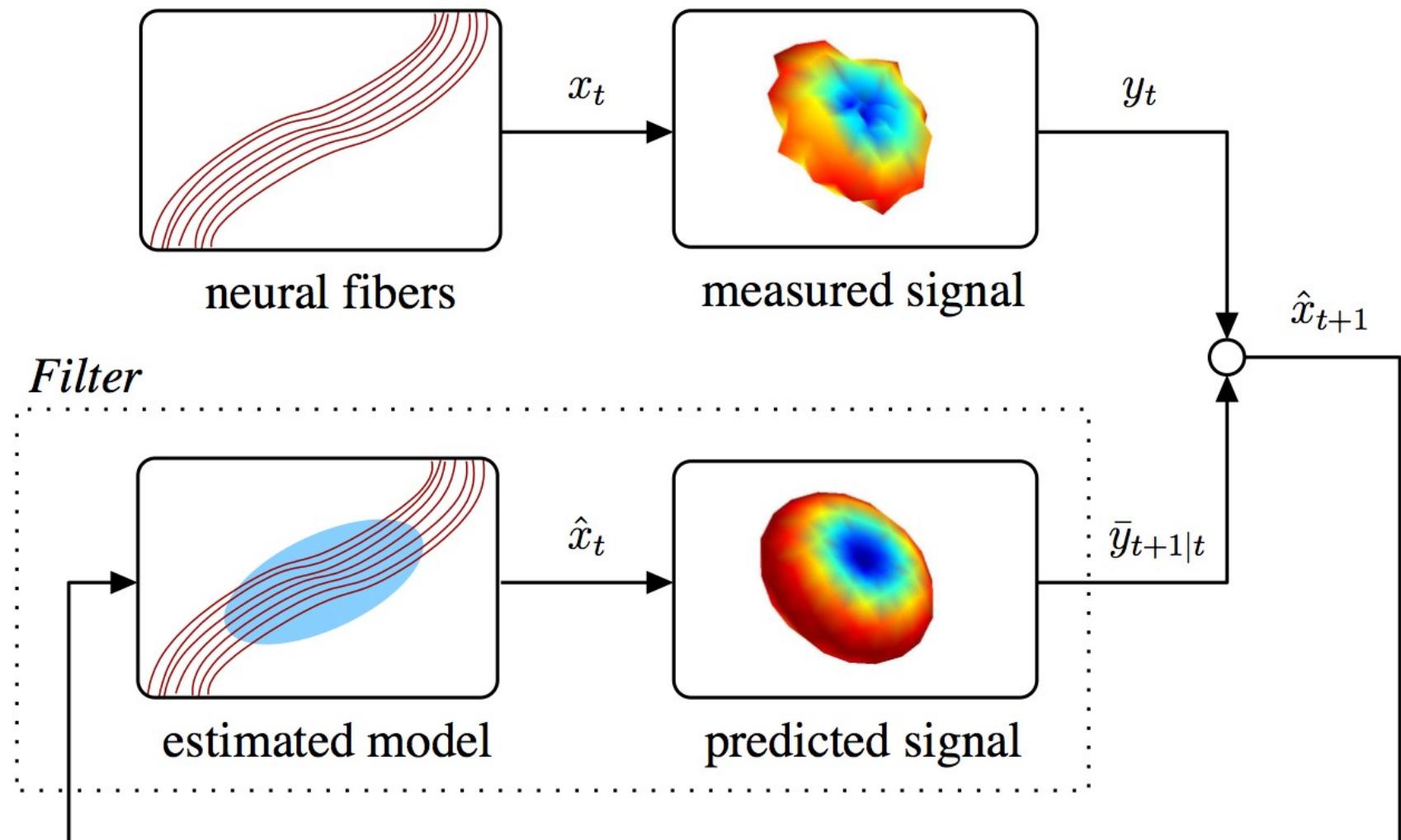
update:  $\boldsymbol{x}_{t+1} = F \boldsymbol{x}_t$      $\boldsymbol{x}_{t+1} = f(\boldsymbol{x}_t)$

observation:  $\boldsymbol{y}_t = G \boldsymbol{x}_t$      $\boldsymbol{y}_t = g(\boldsymbol{x}_t)$

linear

nonlinear

# Kalman filtering



predict ... measure ... reconcile ... repeat ...

$$\begin{aligned} \boldsymbol{x} &= [\boldsymbol{m}_1 \lambda_{11} \lambda_{12} \boldsymbol{m}_2 \lambda_{21} \lambda_{22}]^T \in R^{10} \\ y &\in R^m \text{ signal} \quad \begin{matrix} 10 \text{ dimensional} \\ \text{state} \end{matrix} \end{aligned}$$

$$\boldsymbol{x} = [\boldsymbol{m}_1 \lambda_{11} \lambda_{12} \boldsymbol{m}_2 \lambda_{21} \lambda_{22}]^T \in R^{10}$$

10 dimensional  
state

$$y \in R^m \text{ signal}$$

$$\boldsymbol{x}_{t+1} = f(\boldsymbol{x}_t) = \boldsymbol{x}_t$$

small steps  
slowly varying  
state

$$y_t = g(\boldsymbol{x}_t) = S(\boldsymbol{u})$$

$$\boldsymbol{x} = [\boldsymbol{m}_1 \lambda_{11} \lambda_{12} \boldsymbol{m}_2 \lambda_{21} \lambda_{22}]^T \in R^{10}$$

$$y \in R^m \text{ signal}$$

10 dimensional  
state

$$\boldsymbol{x}_{t+1} = f(\boldsymbol{x}_t) = \boldsymbol{x}_t$$

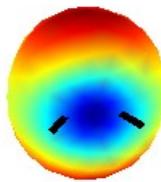
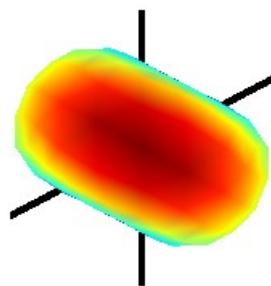
small steps  
slowly varying  
state

$$y_t = g(\boldsymbol{x}_t) = S(\boldsymbol{u})$$

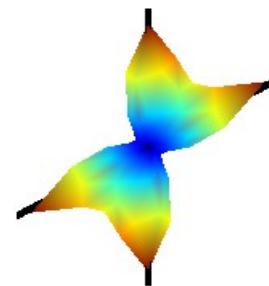
$$y(\boldsymbol{u}) = S(\boldsymbol{u}) = 0.5 s_0 e^{-b\boldsymbol{u}^T D_1 \boldsymbol{u}} + 0.5 s_0 e^{-b\boldsymbol{u}^T D_2 \boldsymbol{u}}$$

$$D = \lambda_1 \boldsymbol{m} \boldsymbol{m}^T + \lambda_2 (\boldsymbol{p} \boldsymbol{p}^T + \boldsymbol{q} \boldsymbol{q}^T)$$

# Local fiber model: mixture of two tensors



diffusion MRI signal



orientation distribution  
function (ODF)

# signal reconstruction is nonlinear

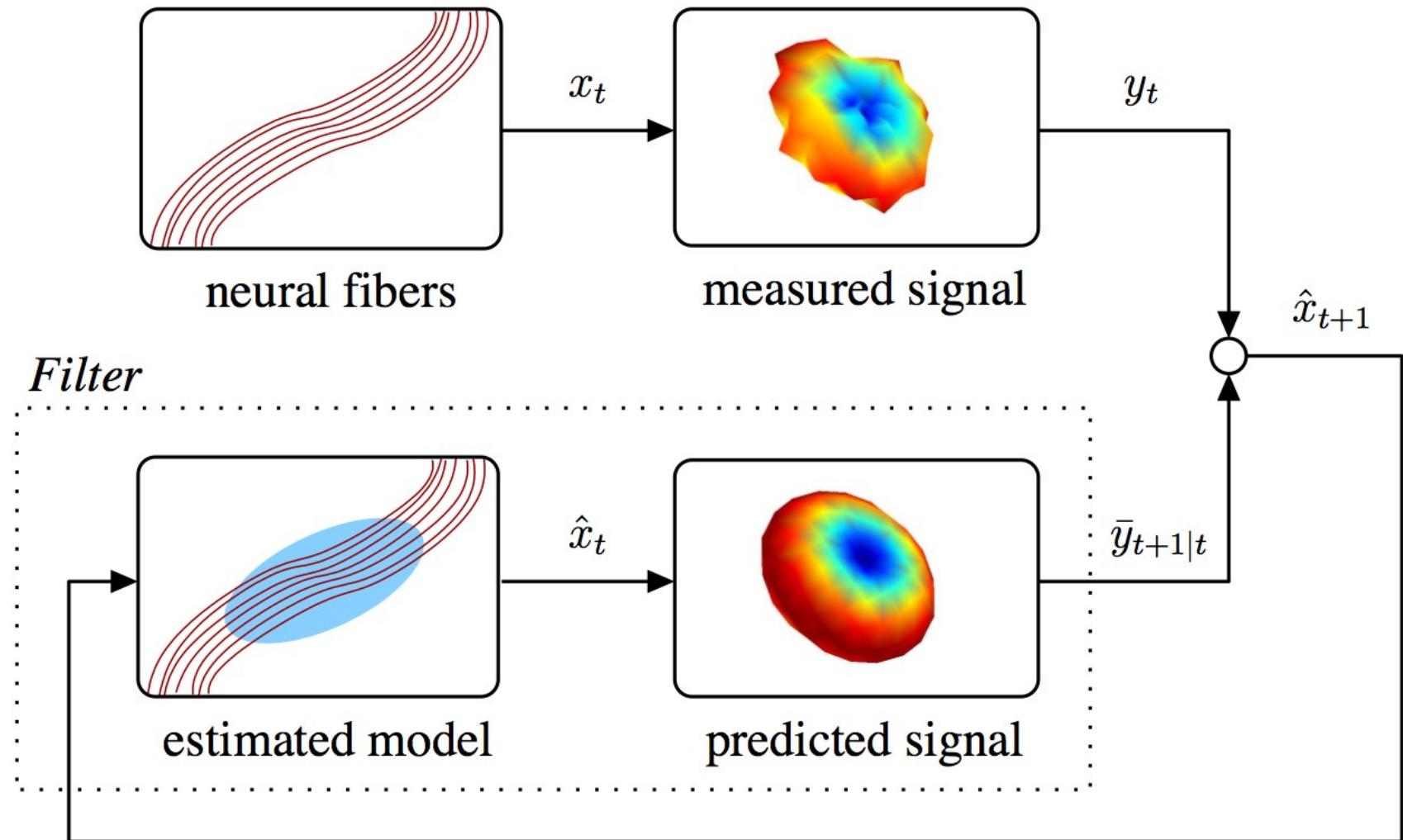
## independent optimization

- least squares  
*linearization*
- gradient descent  
*local minima*
- Levenberg-Marquardt  
*local minima*

## causal estimation

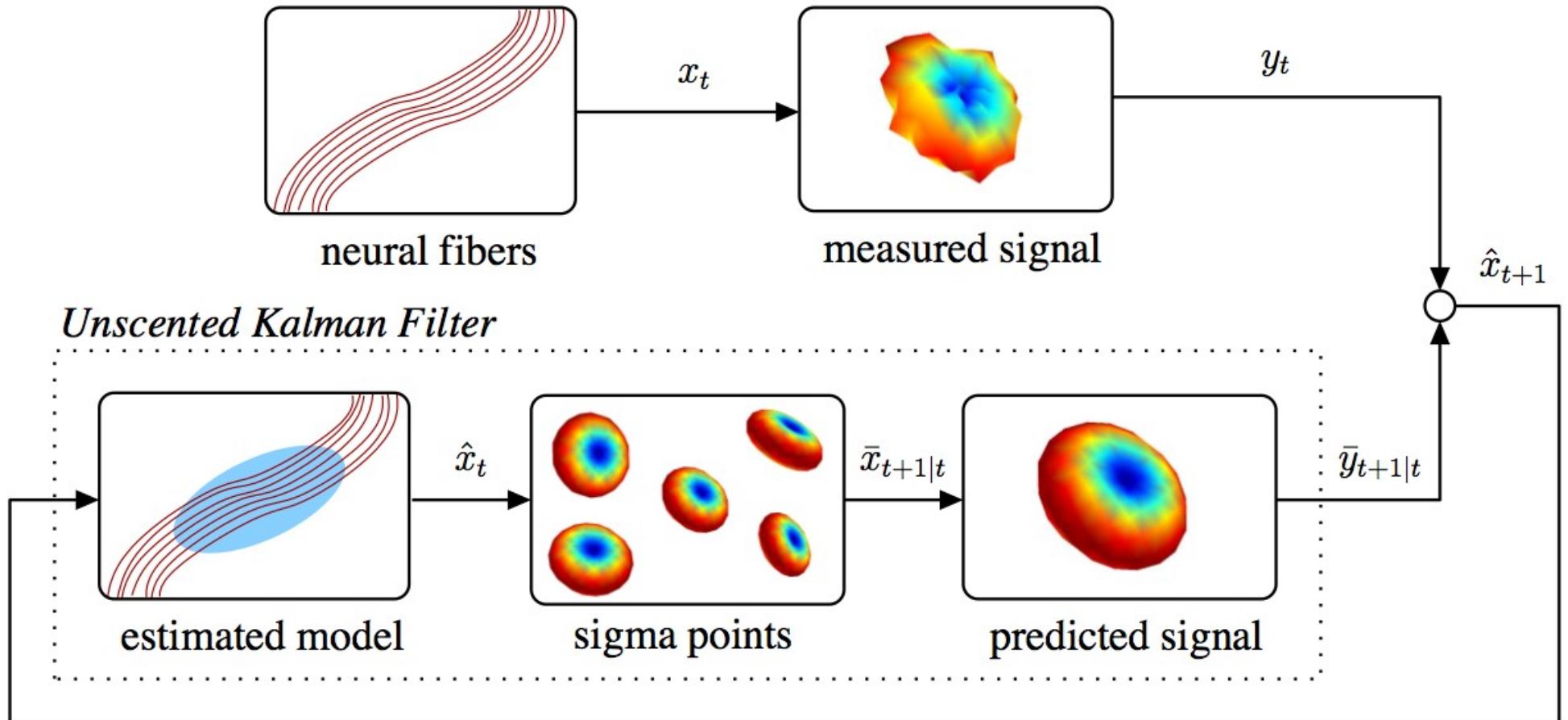
- extended Kalman filter  
mean + covariance  
*linearization*
- particle filter  
non-parametric  
*sampling*
- unscented Kalman filter  
mean + covariance  
no linearization  
limited sampling

# linear Kalman filter



predict ... measure ... reconcile ... repeat ...

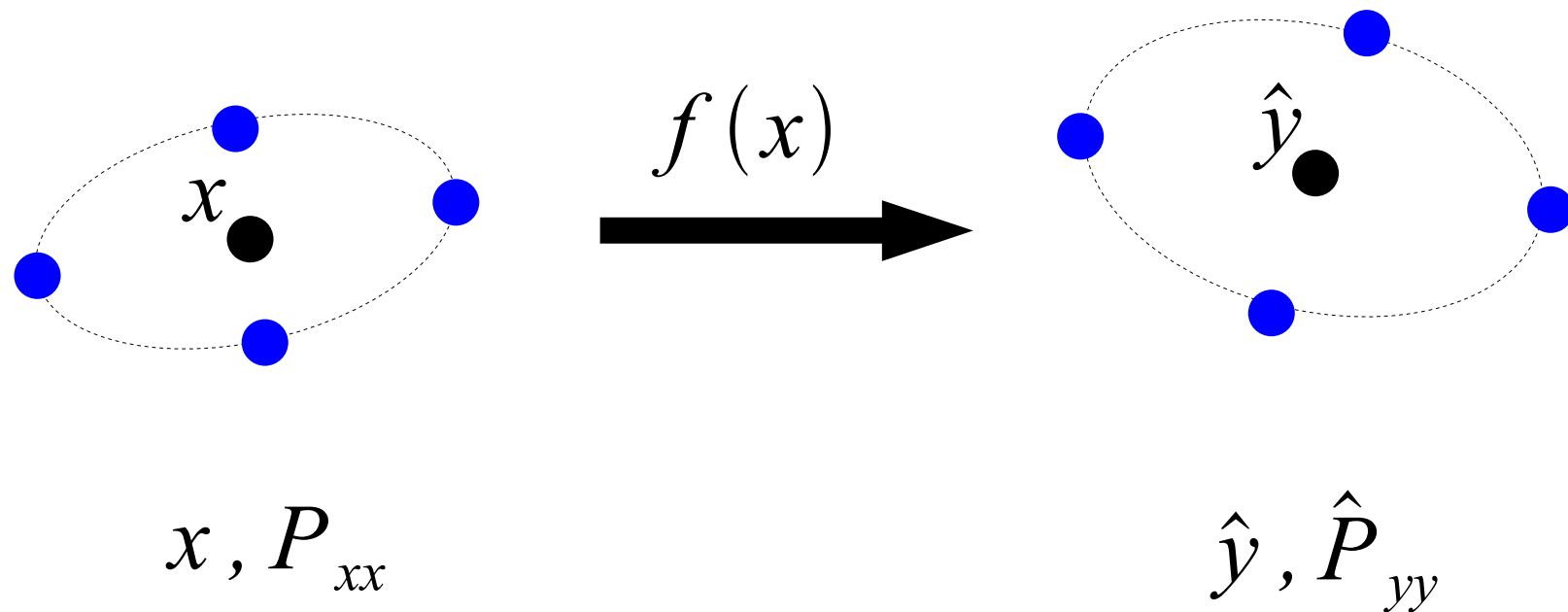
# unscented Kalman filter



same update equations  
modified prediction step

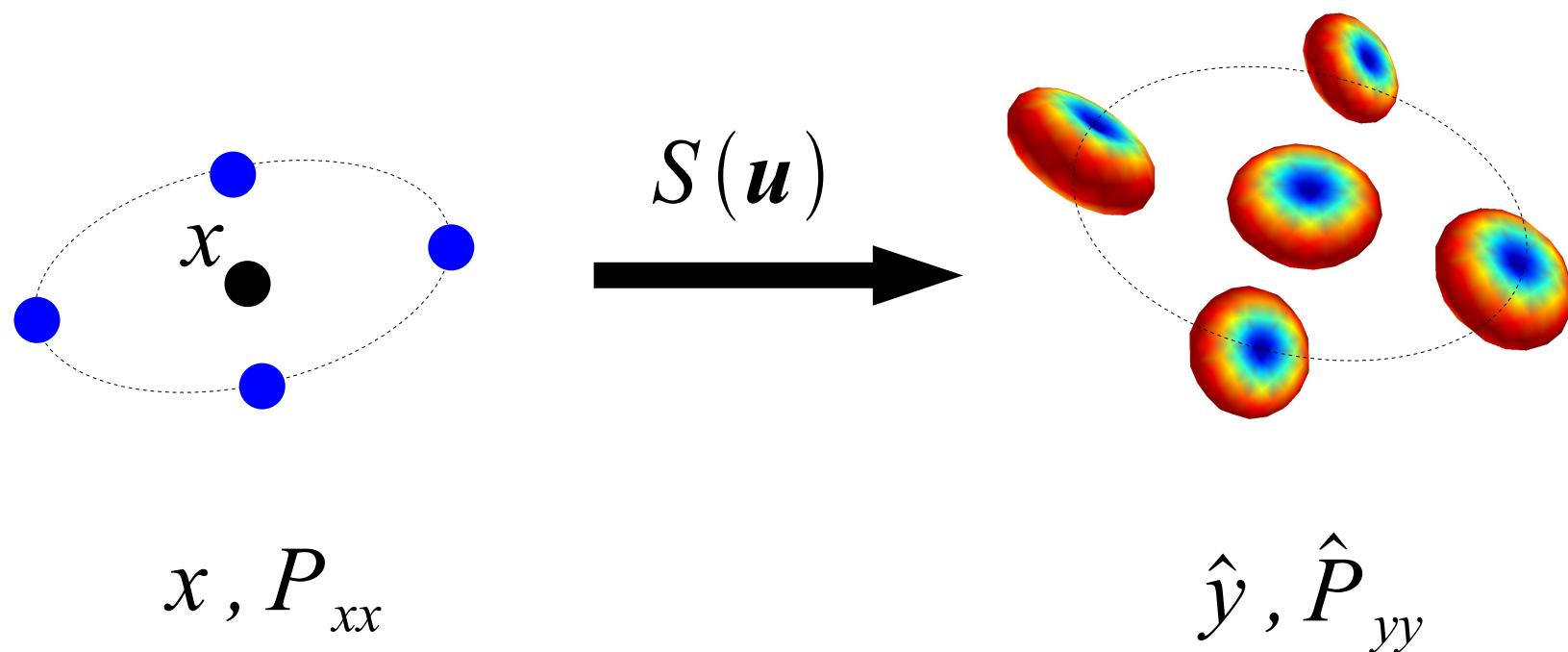
# unscented transform

approximate the statistics...not the function



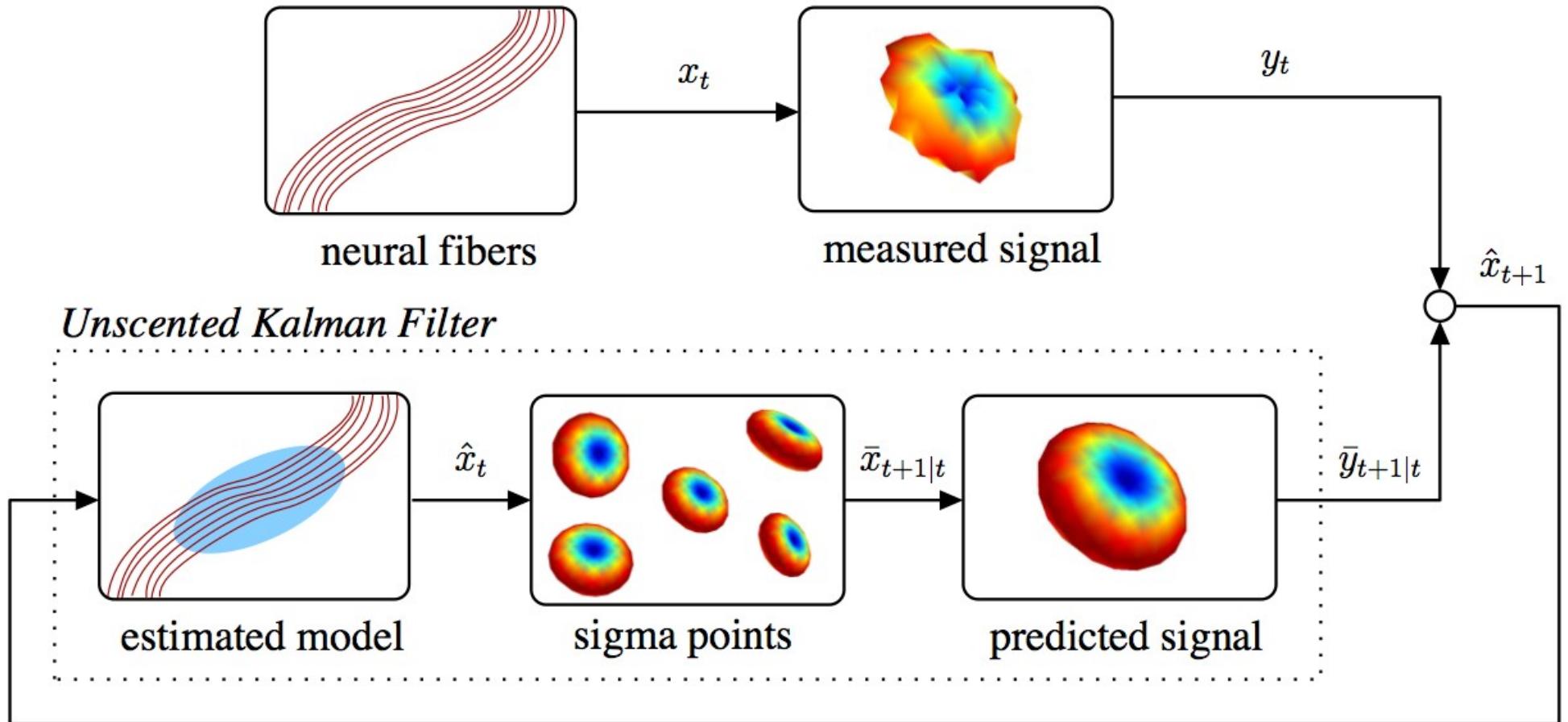
# unscented transform

for signal reconstruction...



$$\boldsymbol{x} = [\boldsymbol{m}_1 k_1 \boldsymbol{m}_2 k_2]^T \in R^8$$

# unscented Kalman filter

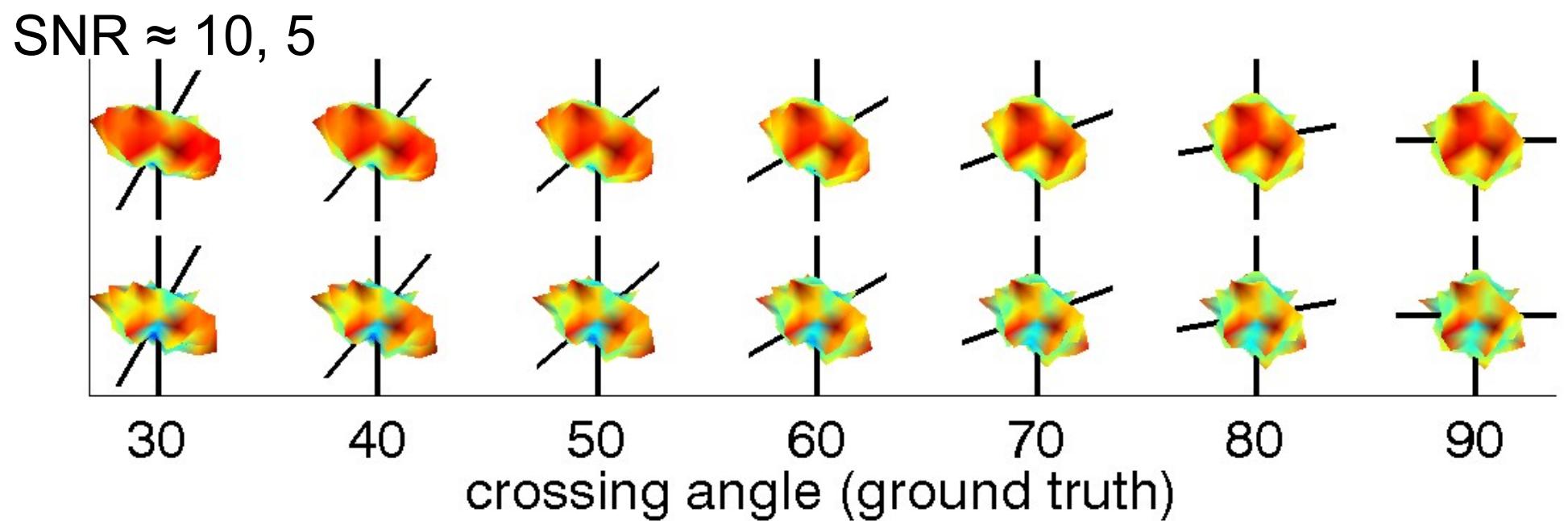
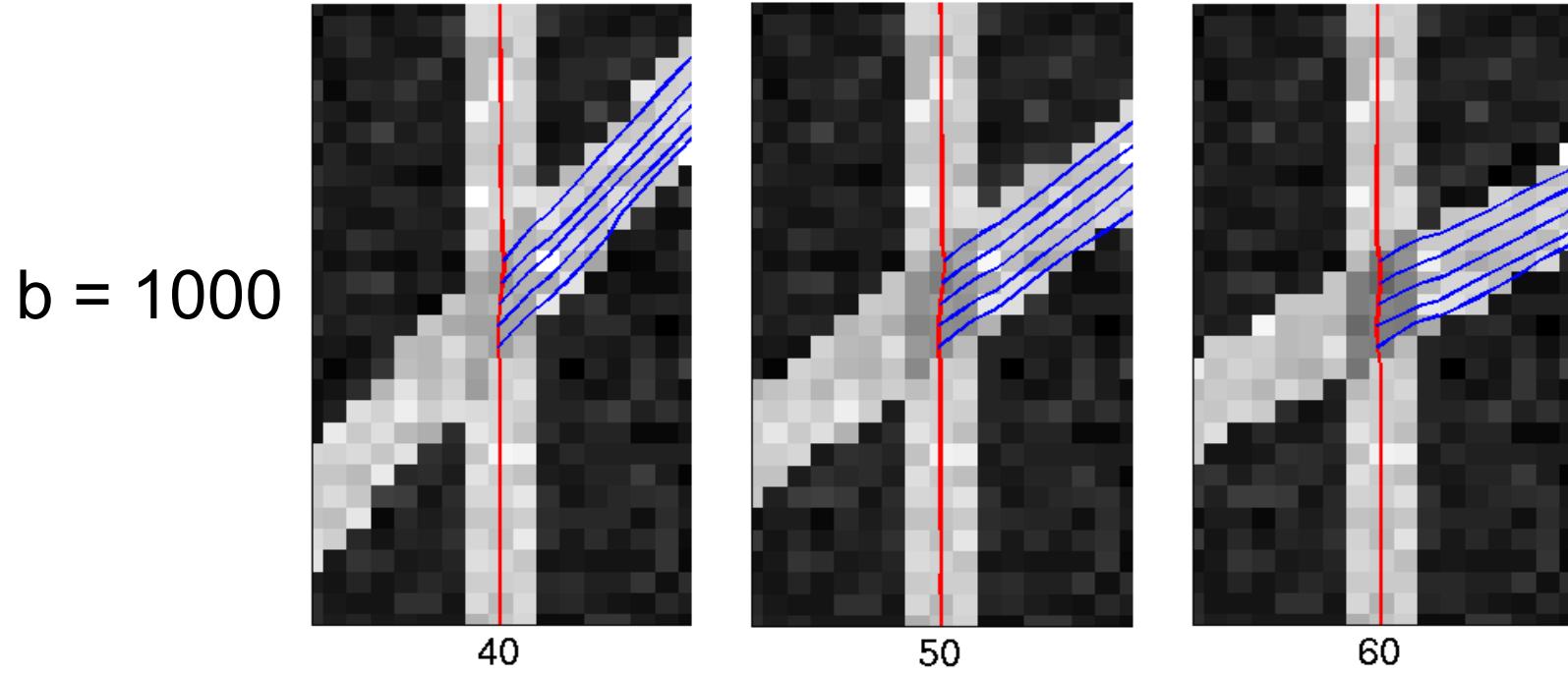


predict ... measure ... reconcile ... repeat ...

# diffusion tensor results

J. G. Malcolm, M. E. Shenton, and Y. Rathi. Filtered multi-tensor tractography. *IEEE Trans. on Medical Imaging*, 29:1664–1675, 2010.

# synthetic validation



## nonlinear least squares

- Levenberg-Marquart
- two-tensor model
- initialize with ground truth (best possible)

## spherical harmonics

- non-parametric
- order eight (8)
- fiber sharpening for peak detection ( $L=0.006$ )

## filtered tractography

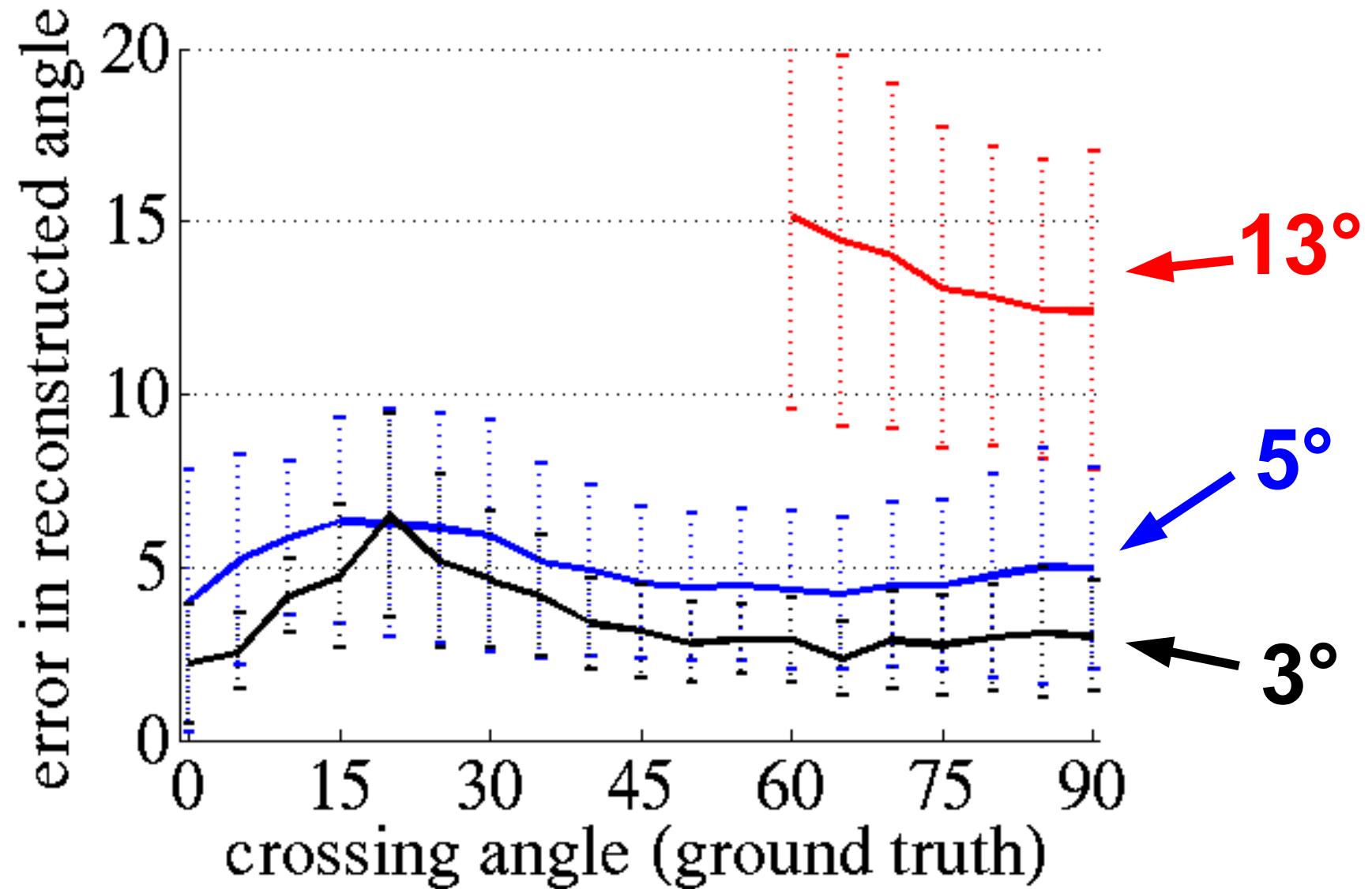
- two-fiber model
- unscented Kalman filter

[Descoteaux 07]

J. G. Malcolm, M. E. Shenton, and Y. Rathi. Neural tractography using an unscented Kalman filter. In *Information Processing in Medical Imaging (IPMI)*, pages 126–138, 2009.

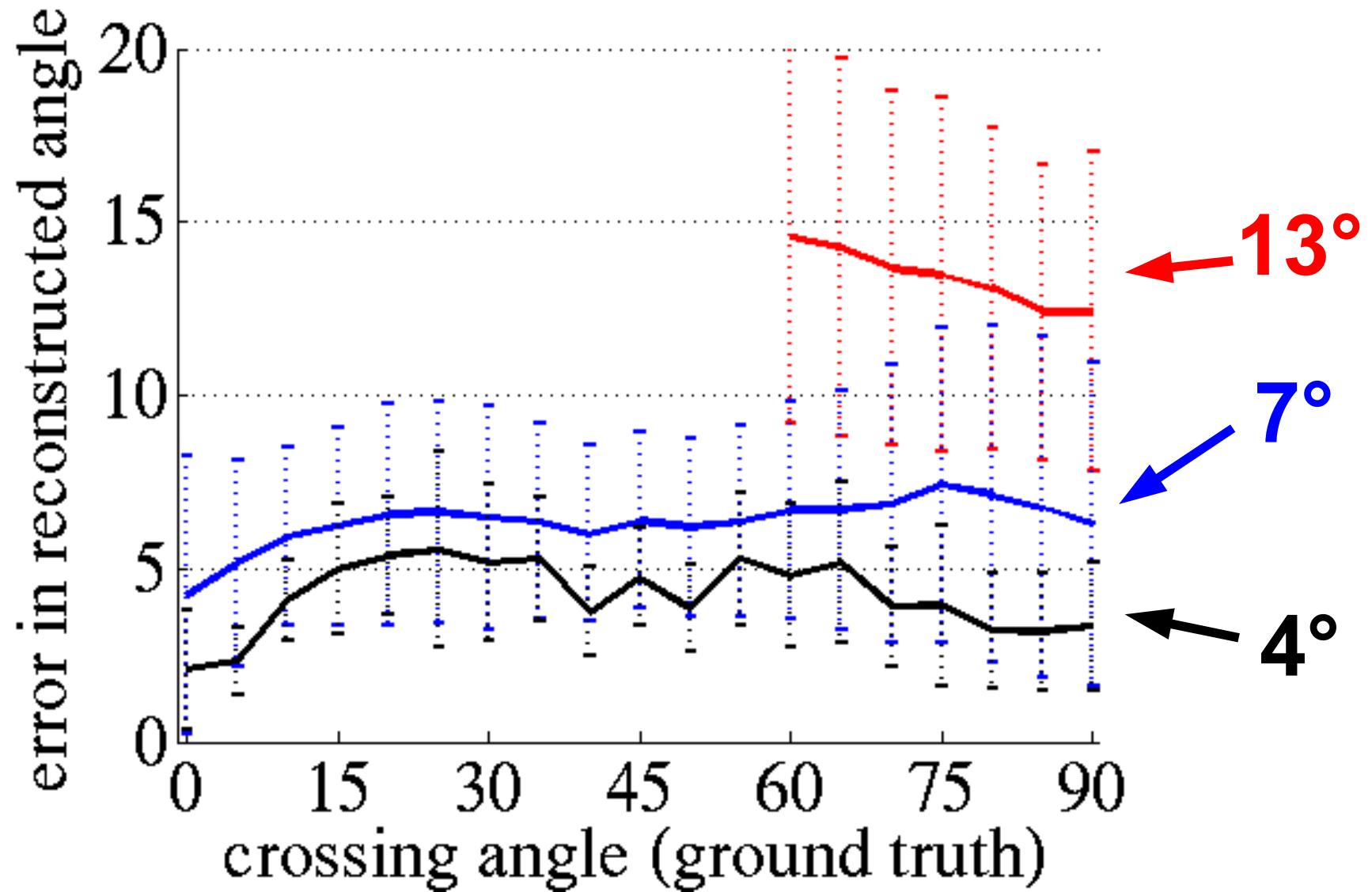
J. G. Malcolm, M. E. Shenton, and Y. Rathi. Filtered multi-tensor tractography. *IEEE Trans. on Medical Imaging*, 29:1664–1675, 2010.

# angular reconstruction error



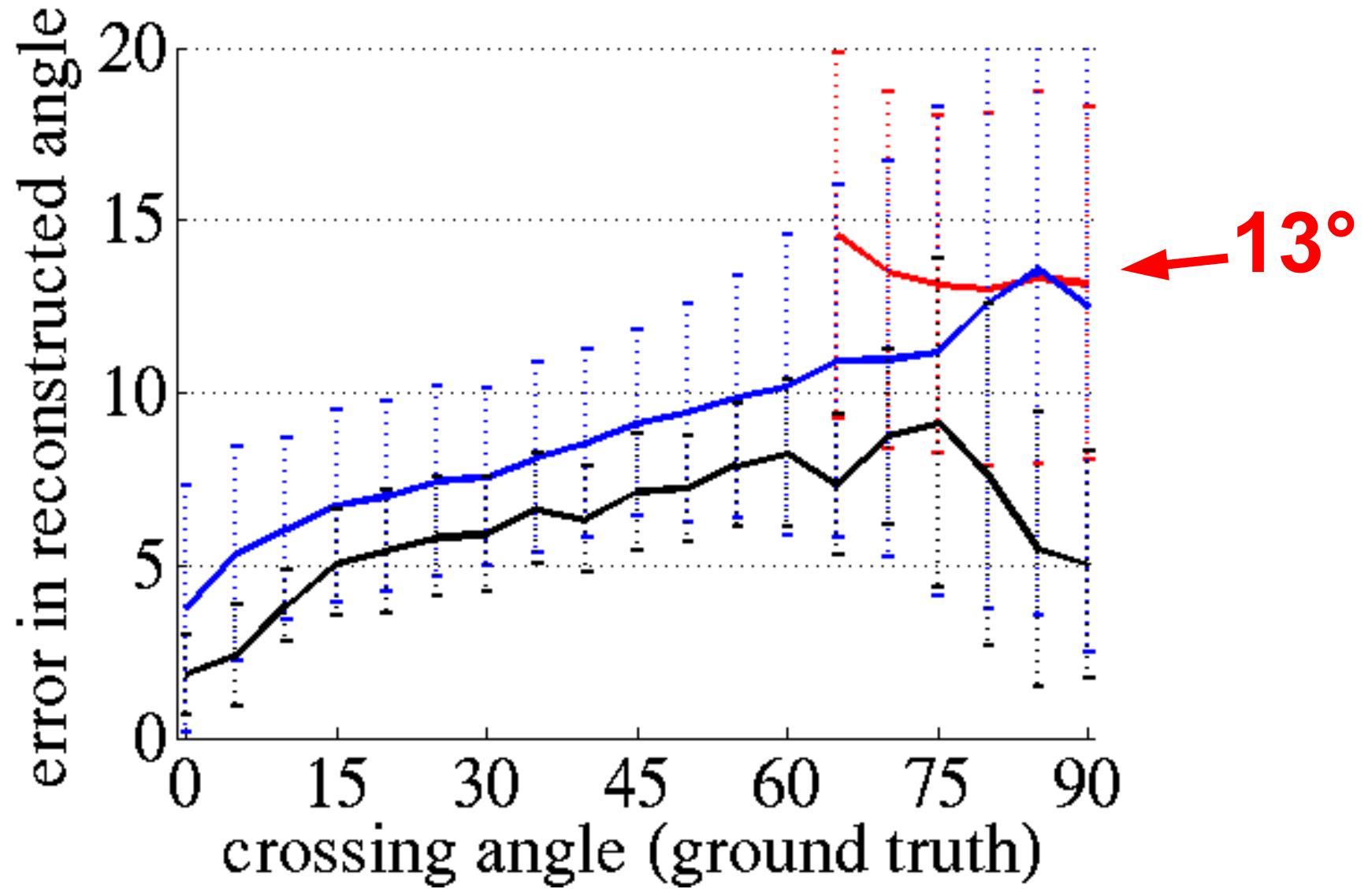
SNR  $\approx 5$ , b = 1000

# unequal signal field: 60/40



SNR  $\approx 5$ , b = 1000

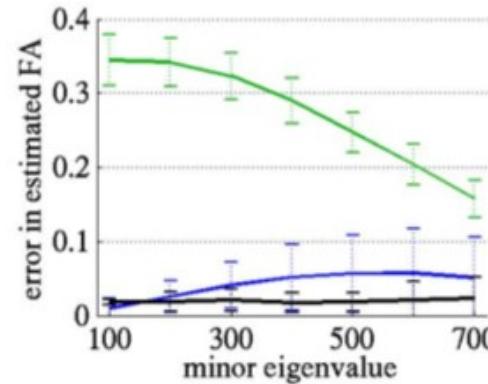
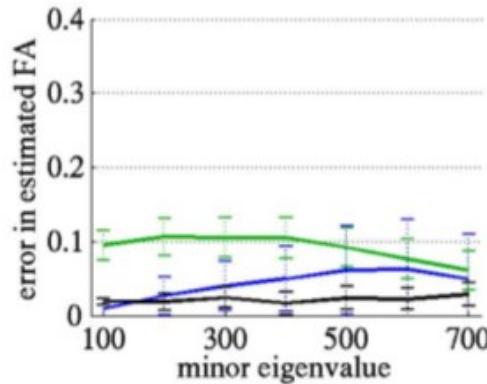
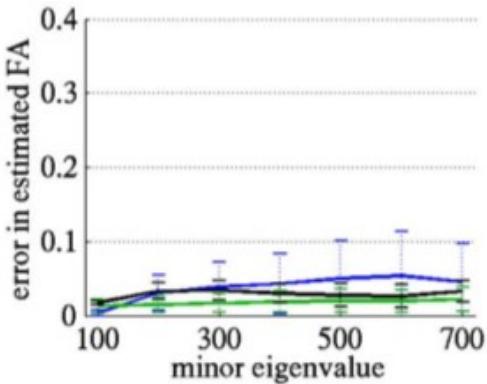
# unequal signal field: 70/30



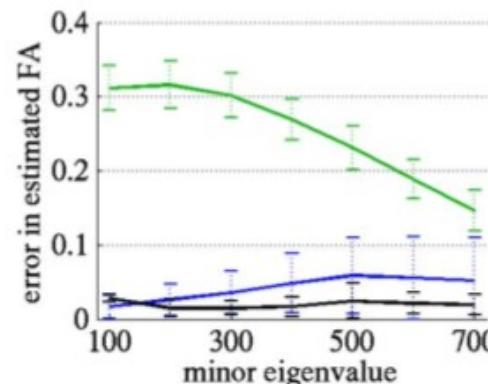
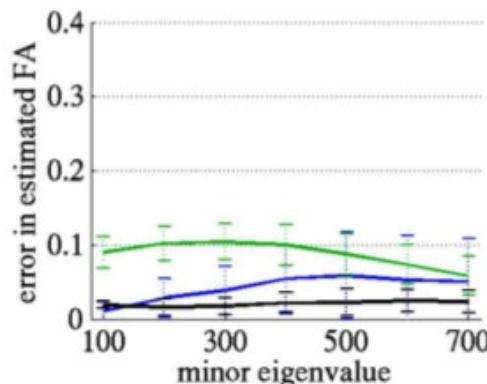
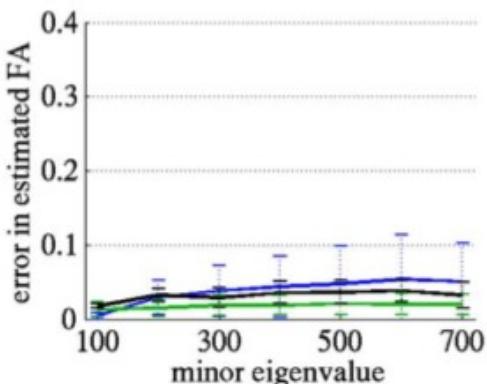
SNR  $\approx 5$ , b = 1000

# estimation fidelity

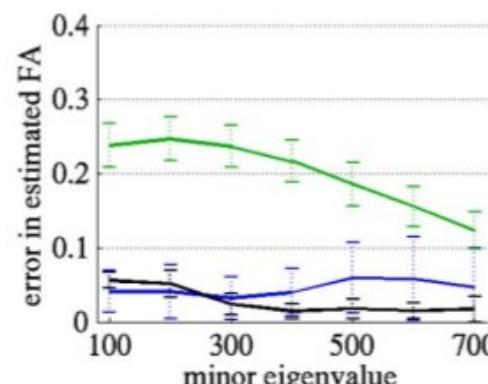
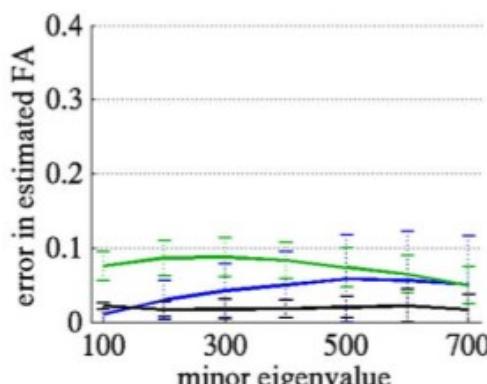
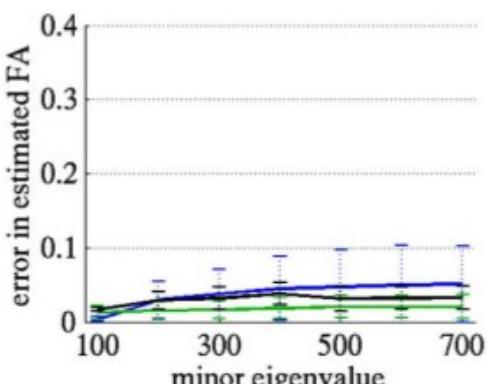
50/50



60/40



70/30



one fiber  
(no crossing)

45 degree crossing

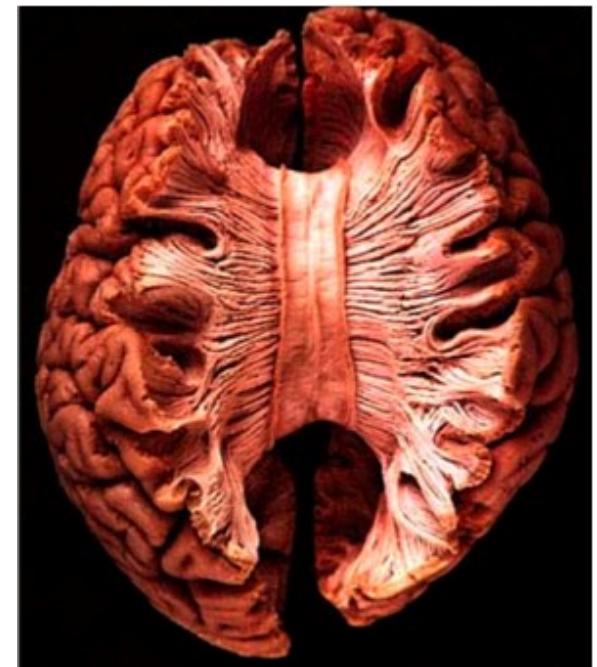
90 degree crossing



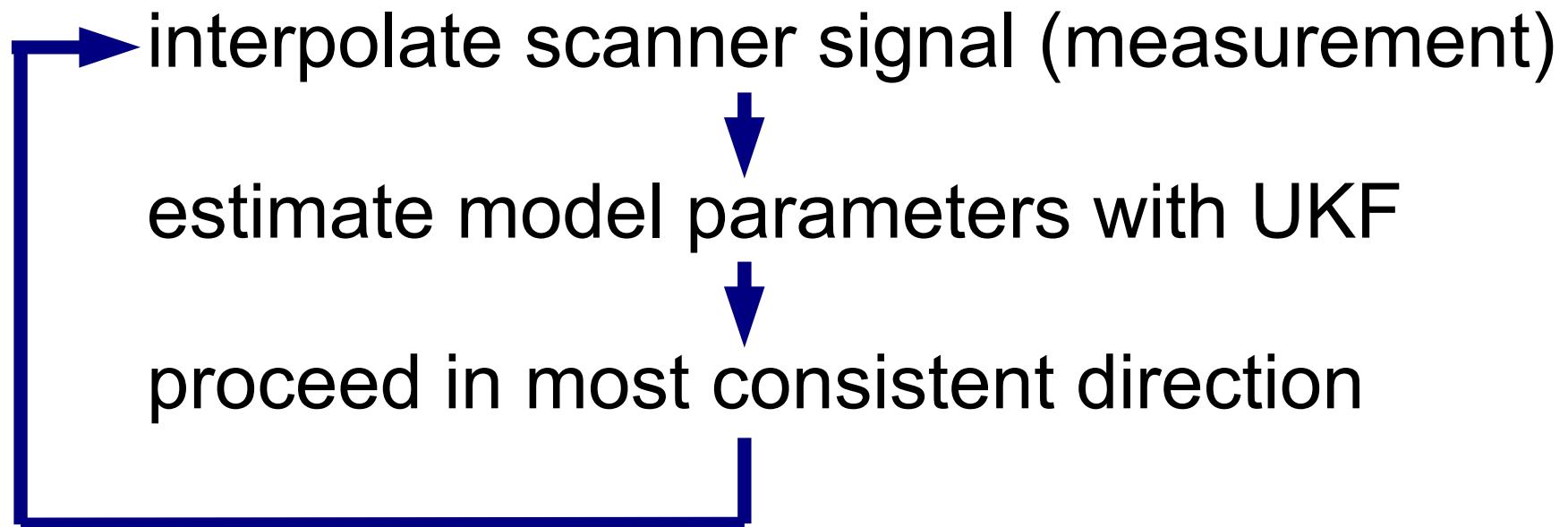
in vivo



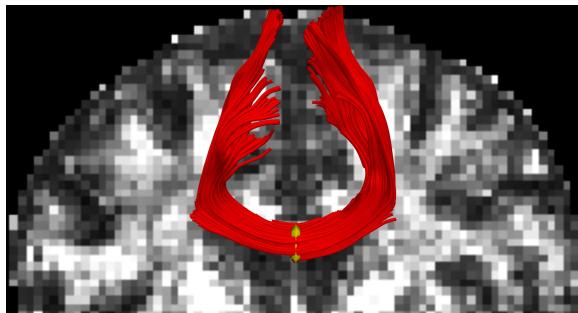
b=900  
51 directions  
1.7mm isotropic voxel  
17 minute scan



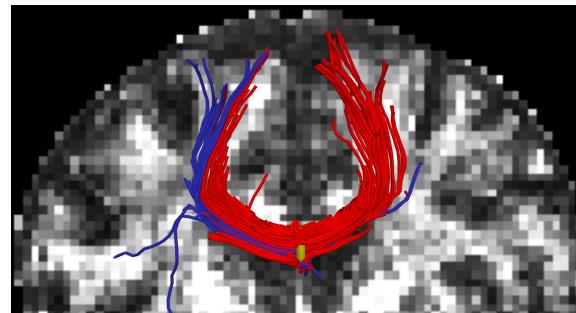
# algorithm



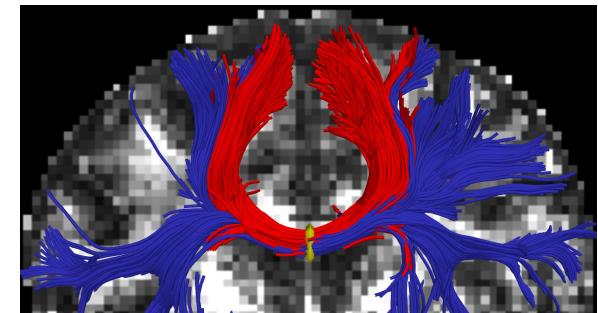
terminate:  $FA < 0.15$



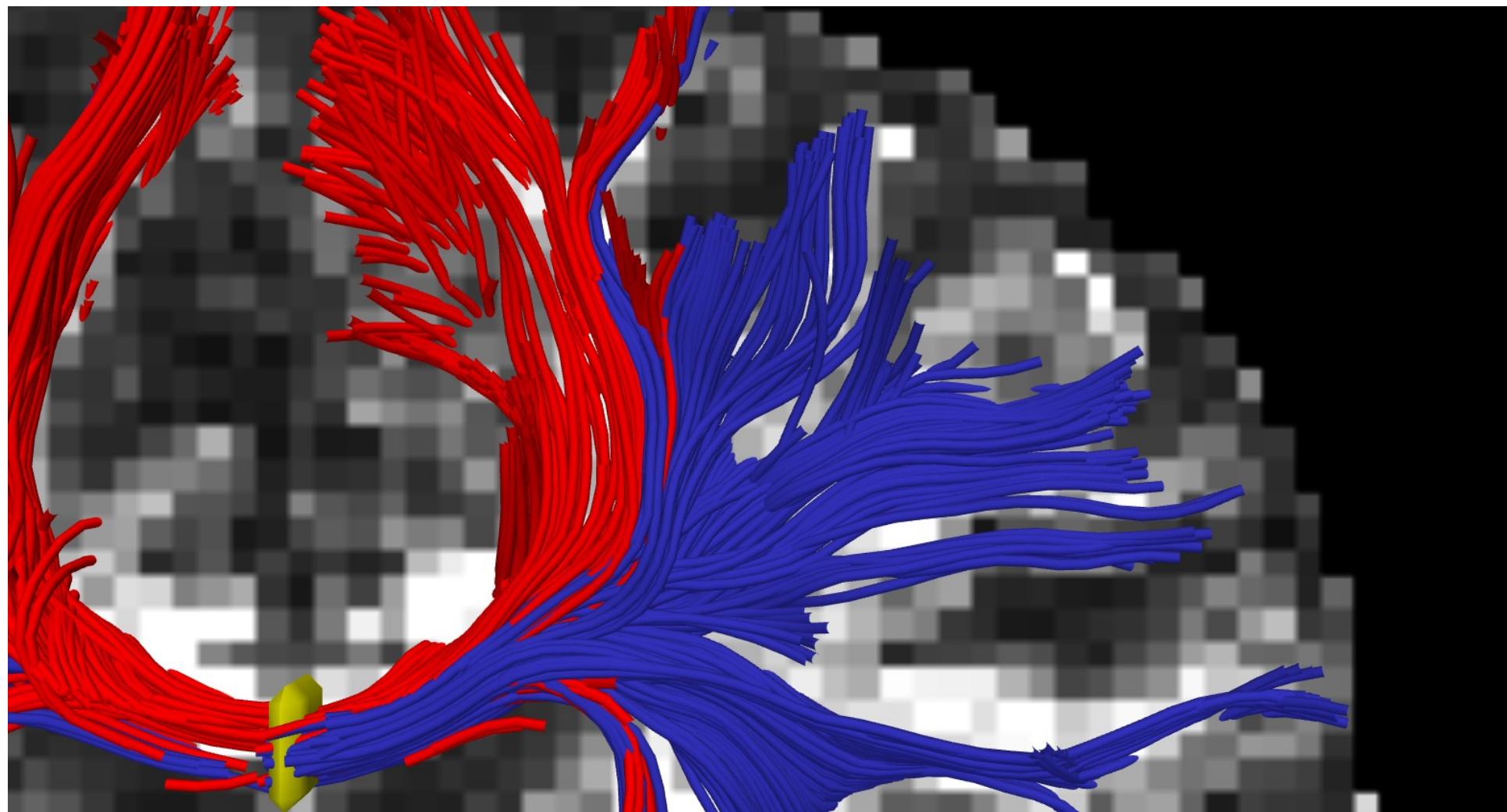
single tensor



spherical harmonics



filtered two-tensor

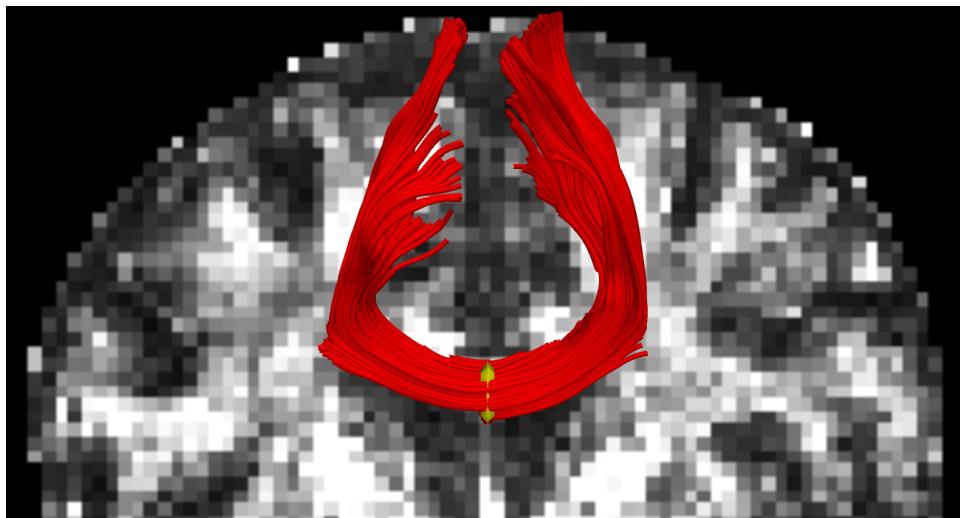


filtered two-tensor

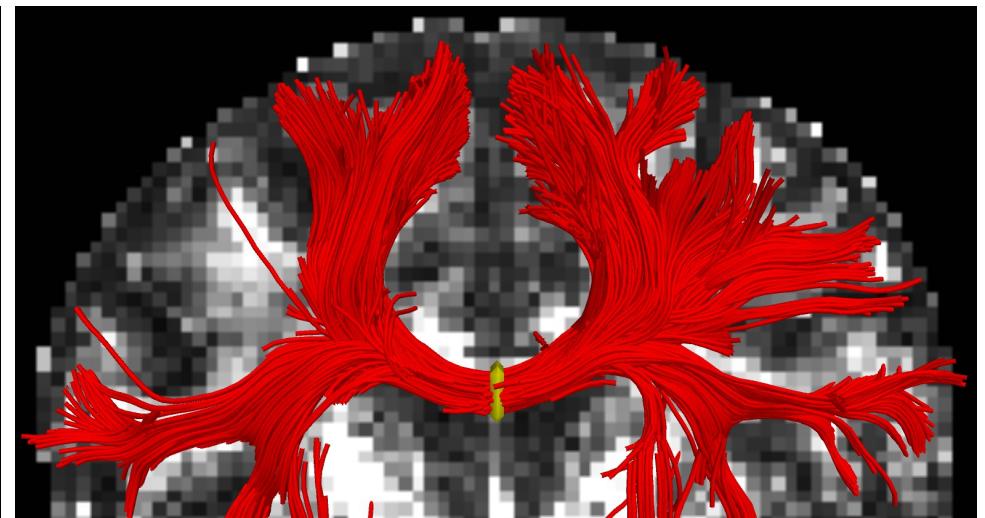
( $b = 900$ , 1.7mm, 51 directions)

# fitting the local fiber model

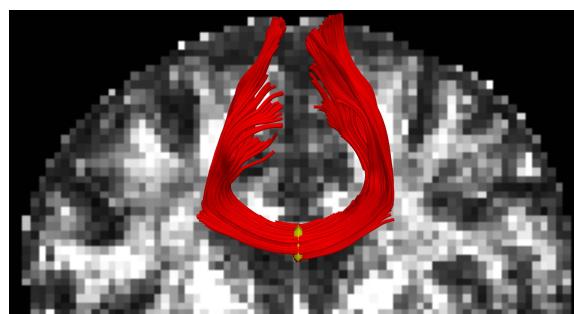
- Independent optimization v. filtered estimation
- Nonlinear model: Unscented Kalman filter



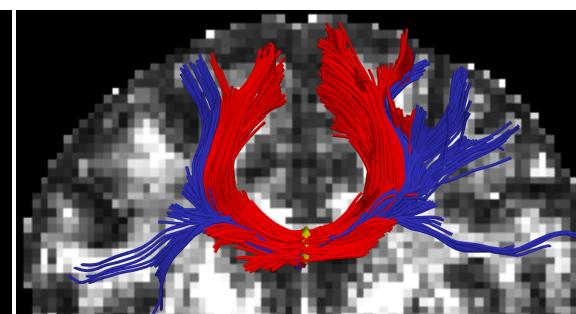
single-tensor



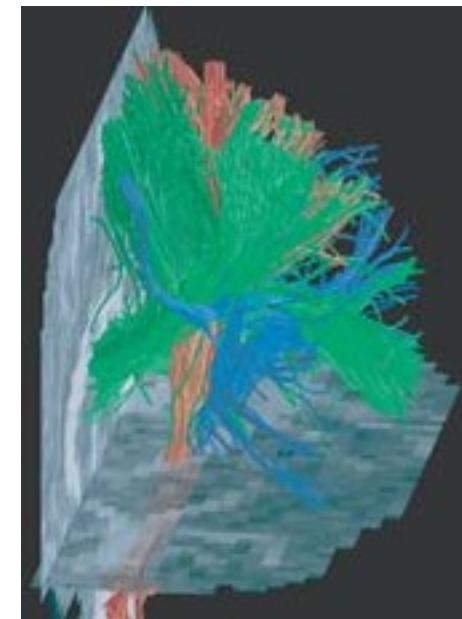
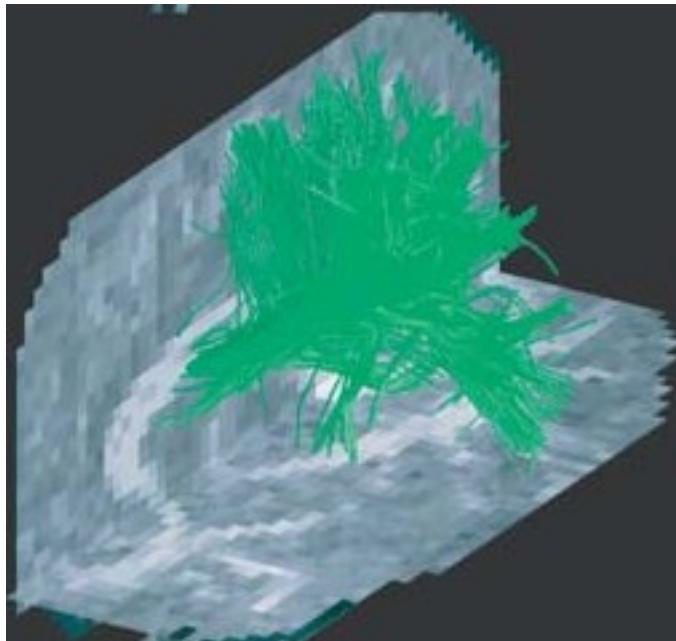
filtered two-tensor



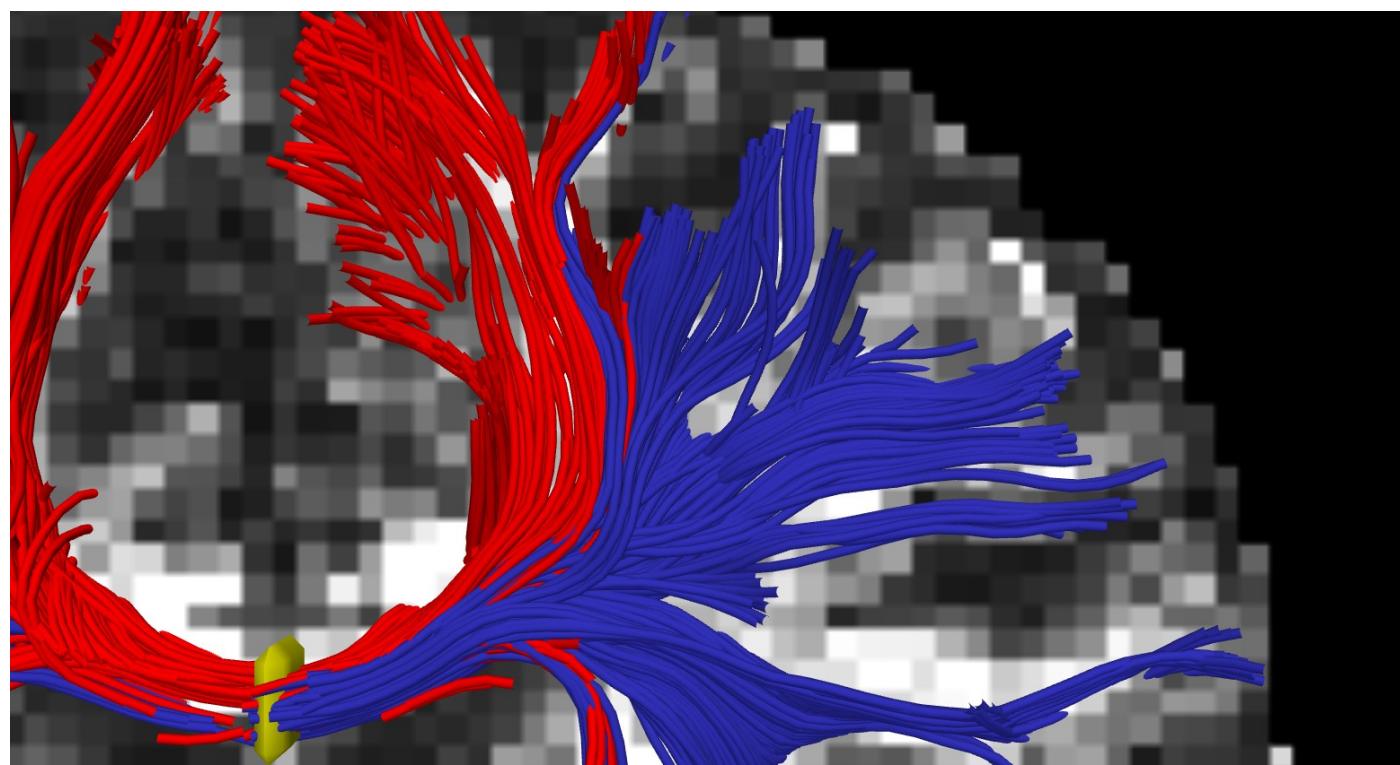
single tensor



filtered two-tensor

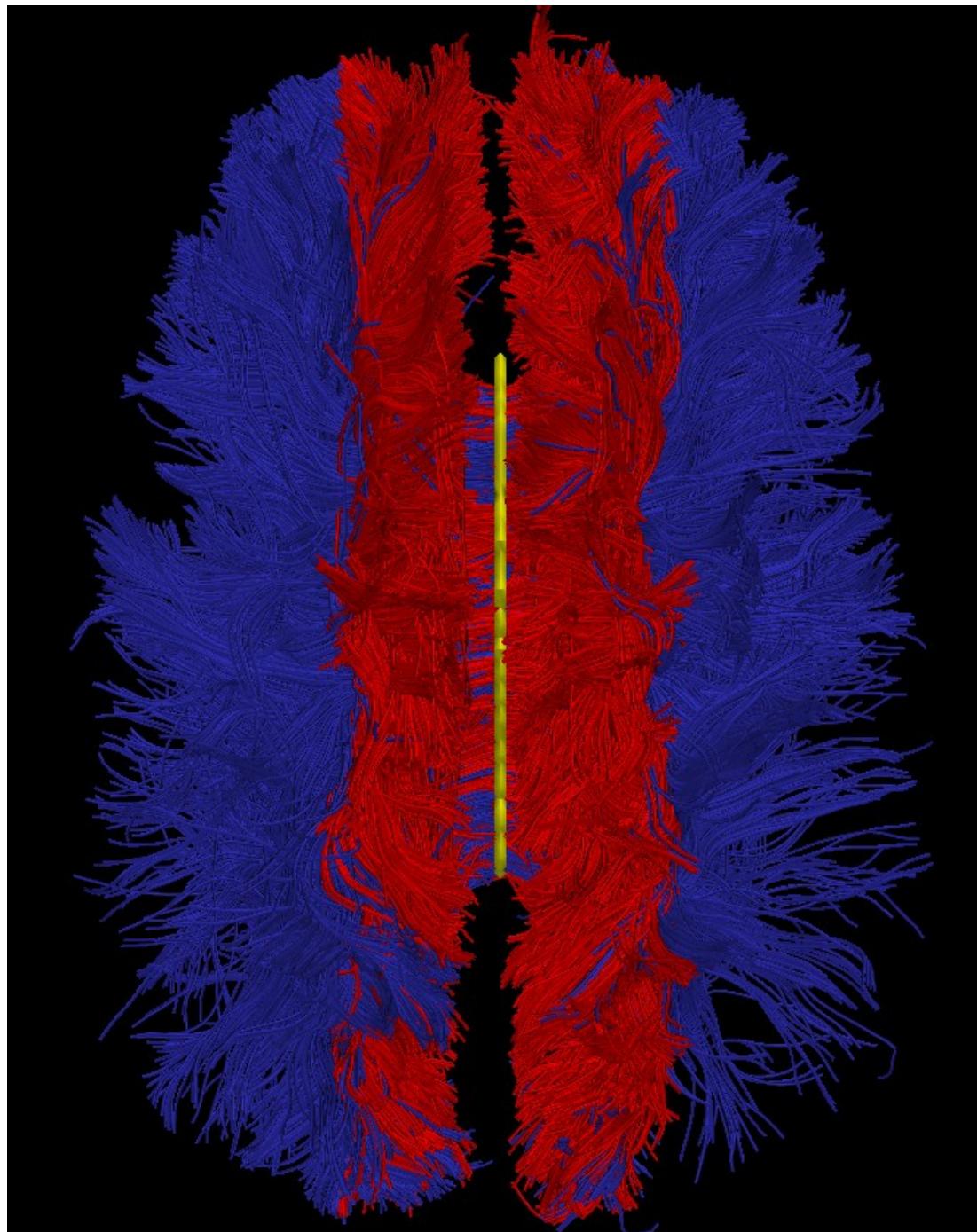


DSI [Hagmann 05]



filtered two-tensor

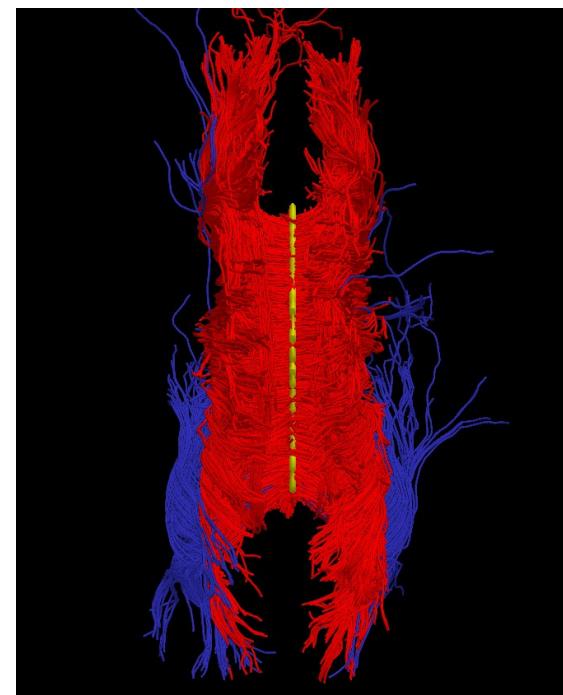




filtered two-tensor



single tensor

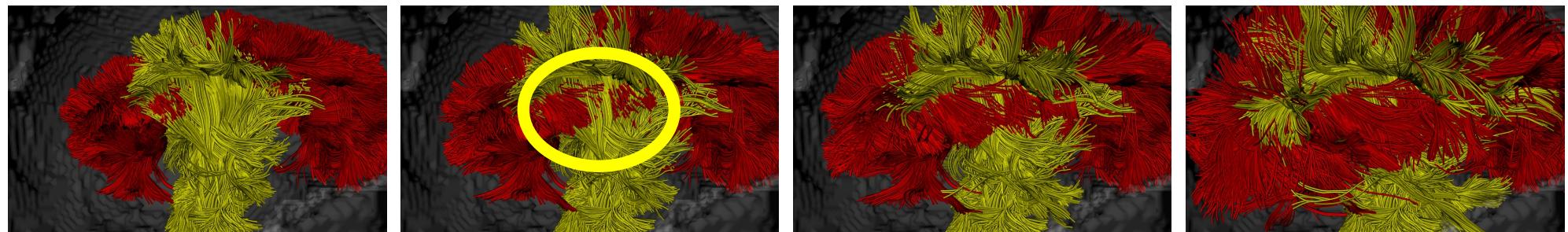


spherical harmonics

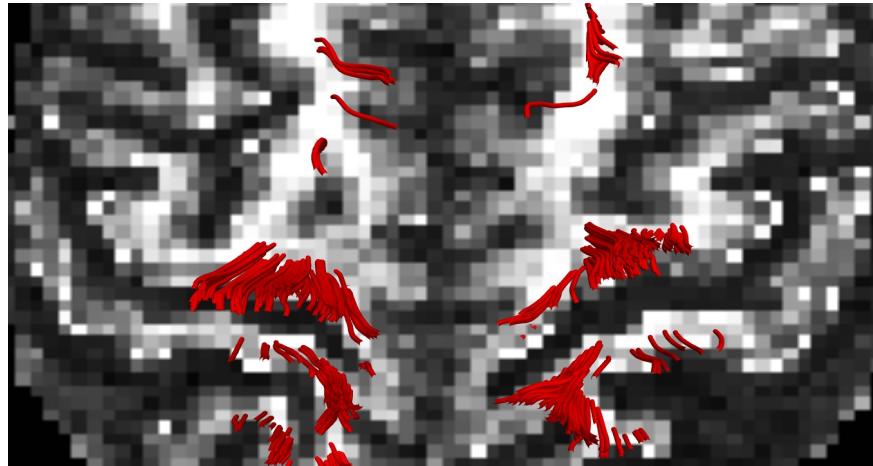


corpus callosum

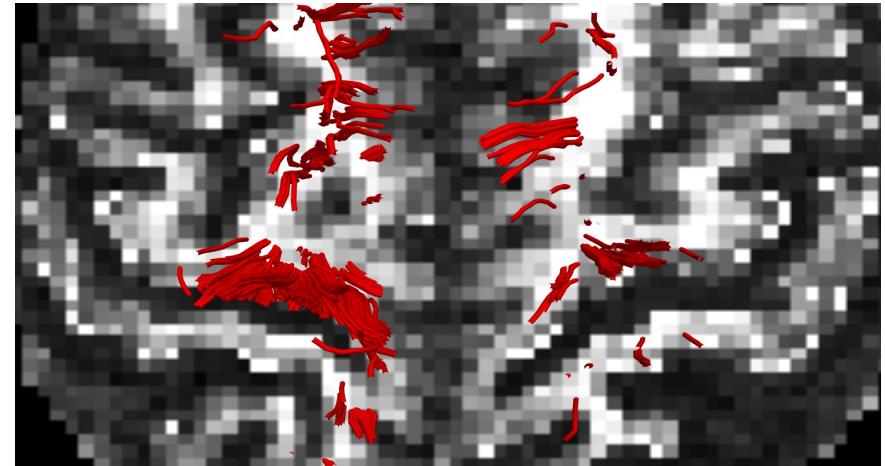
internal capsule



centrum semiovale



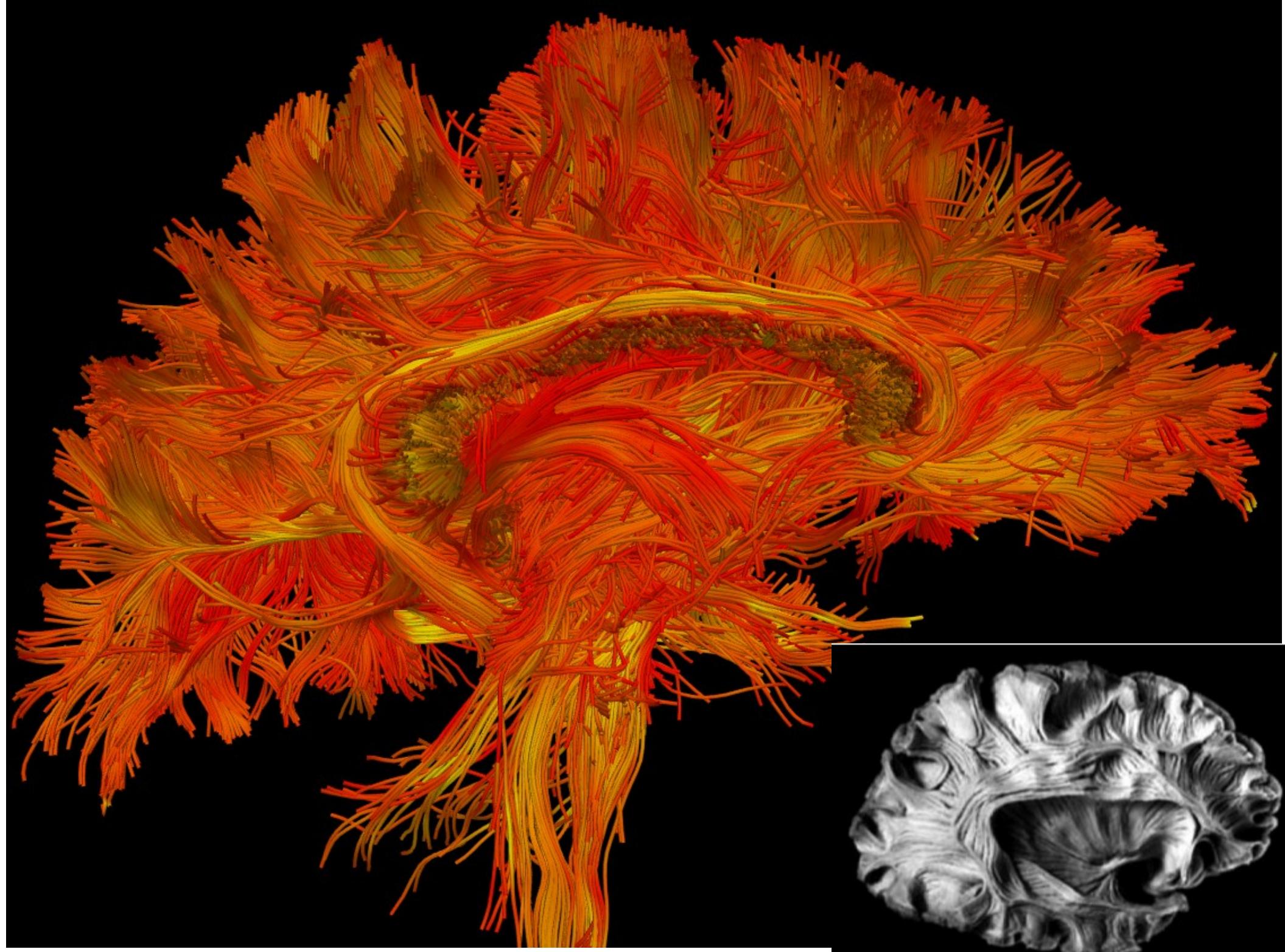
single tensor



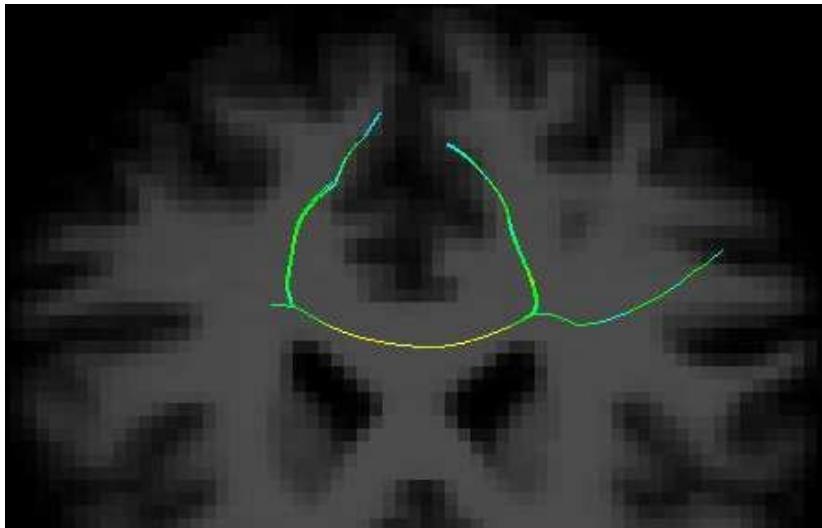
spherical harmonics



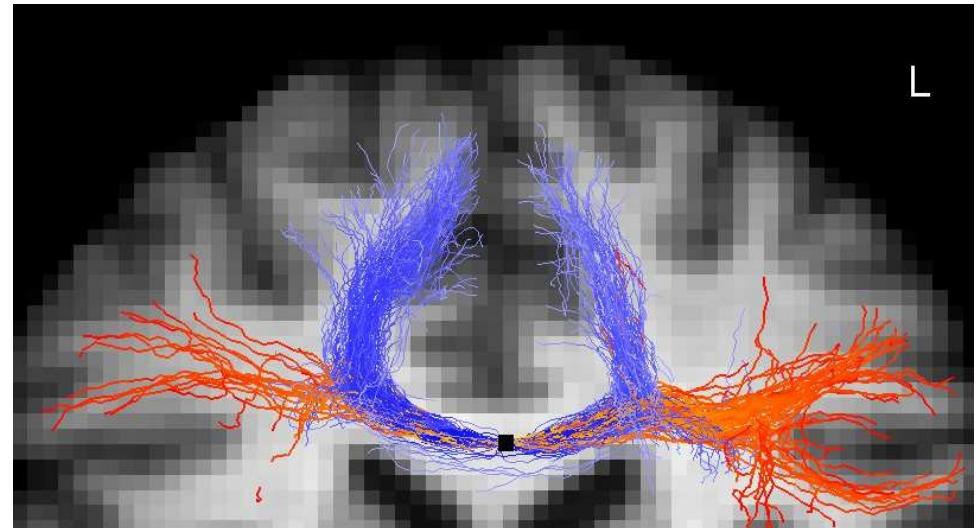
filtered two-tensor



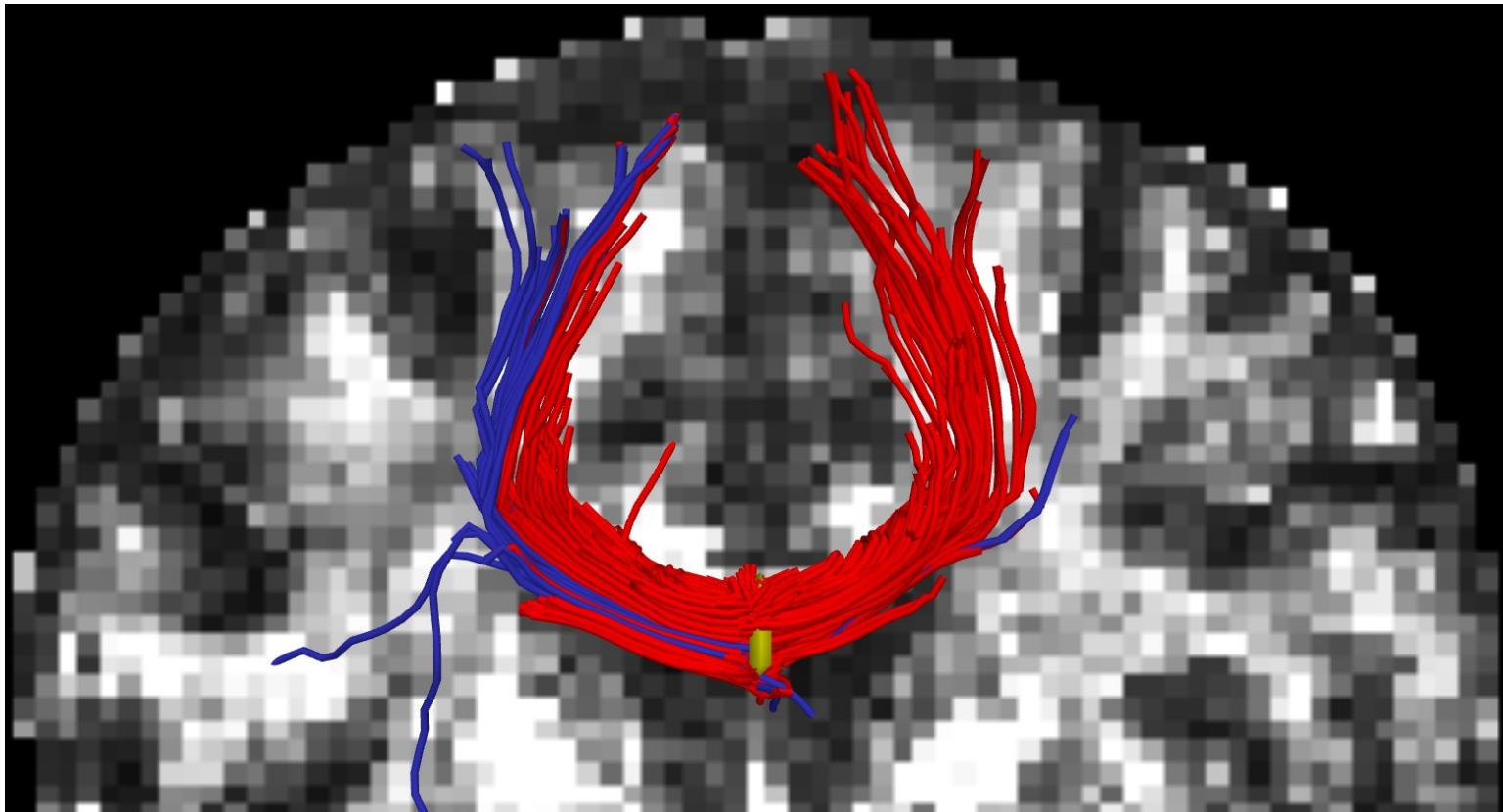
# related work



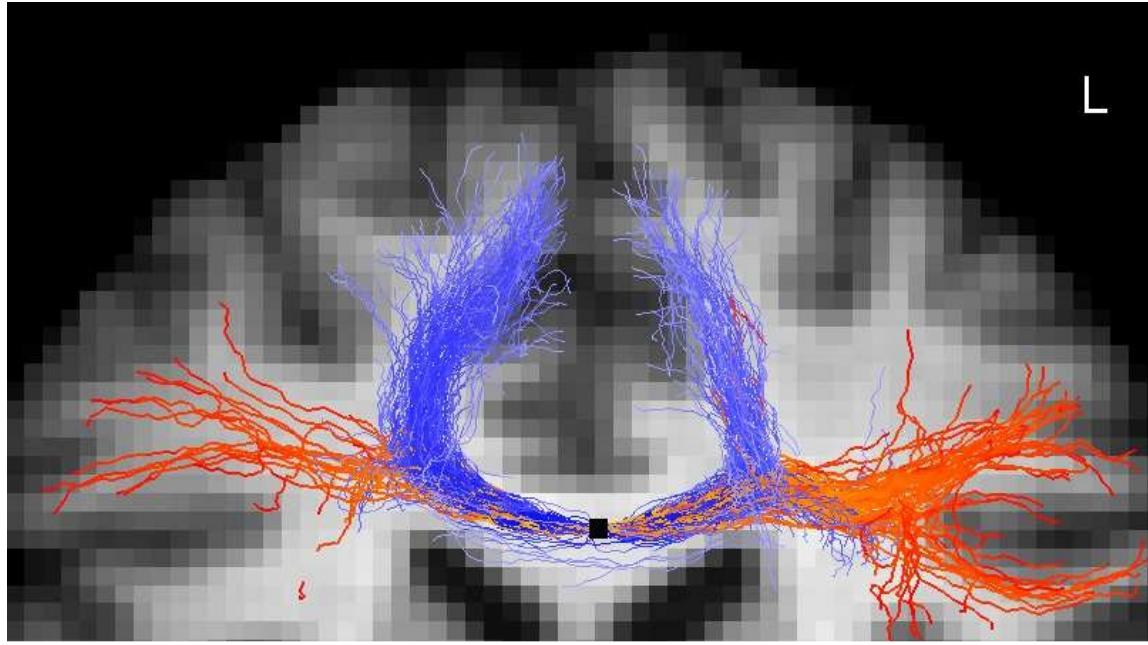
Deterministic (Descoteaux 2007)



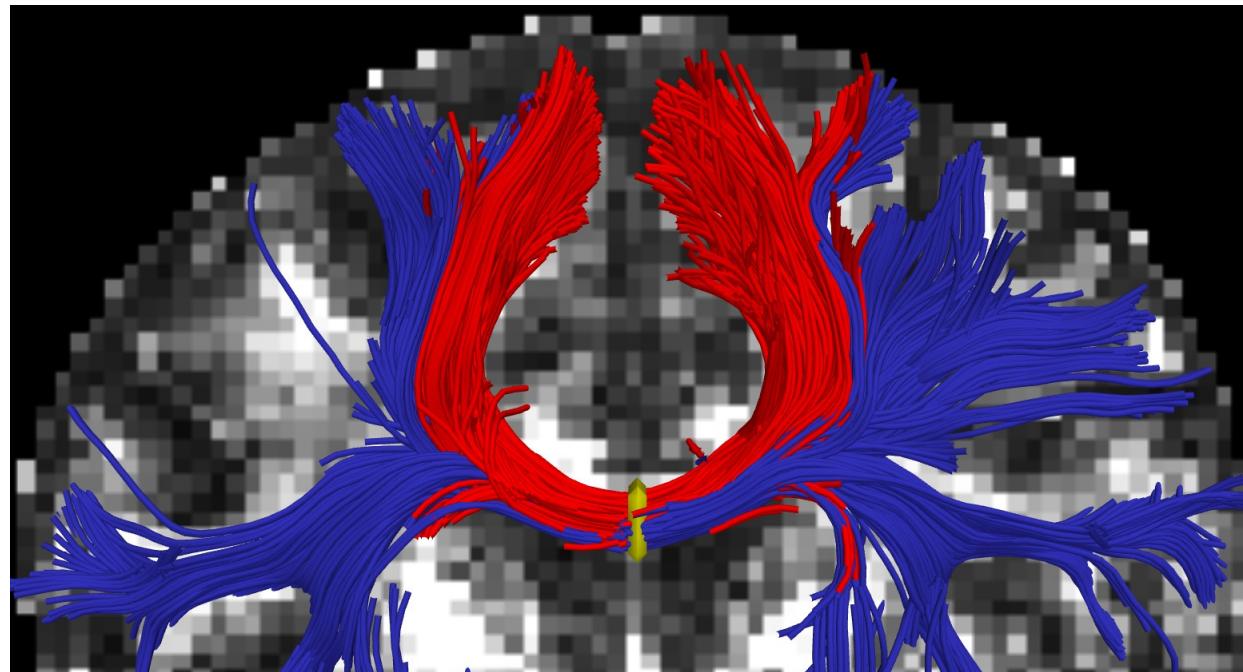
Probabilistic (Descoteaux 2007)



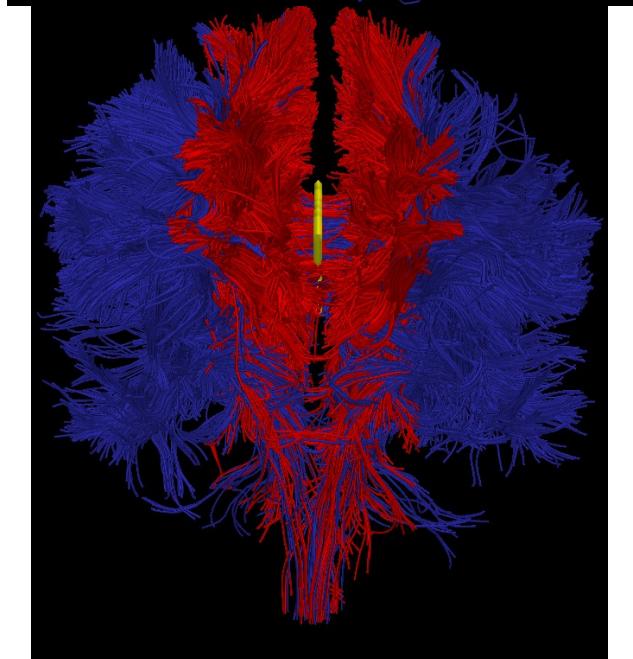
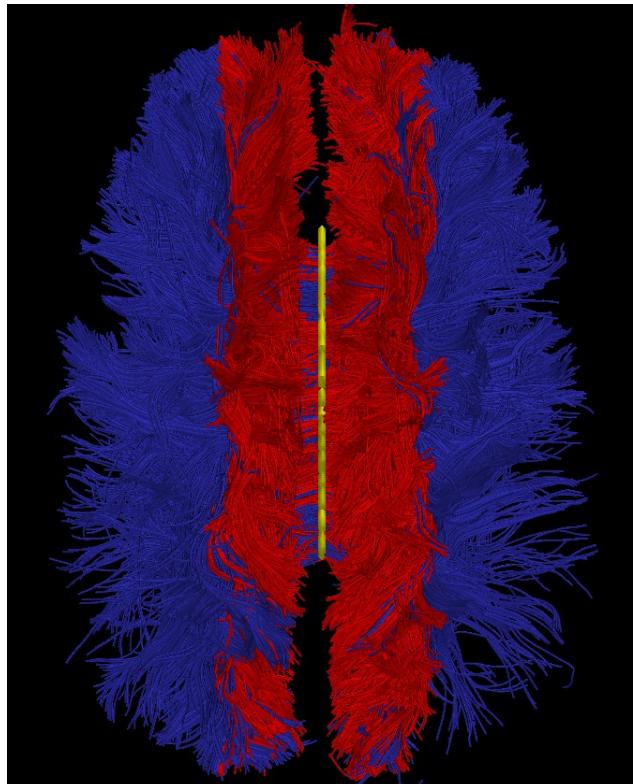
our implementation of spherical harmonics



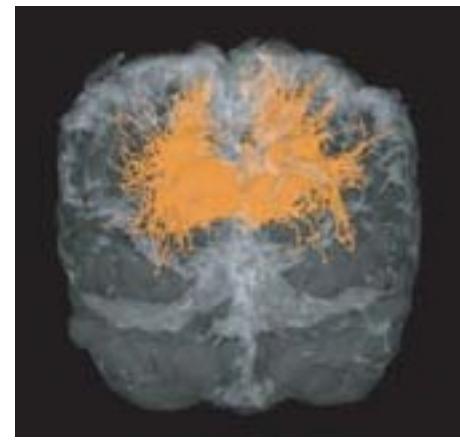
spherical harmonics, **probabilistic** tractography  
[Descoteaux 2007]



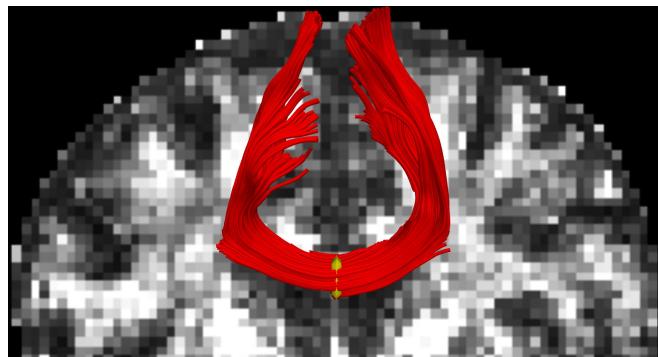
filtered two-tensor, **deterministic** tractography



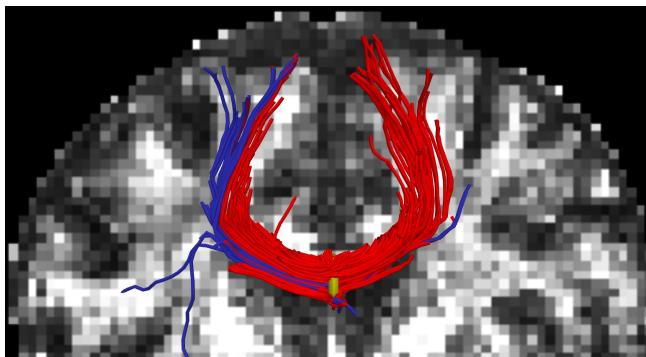
filtered two-tensor



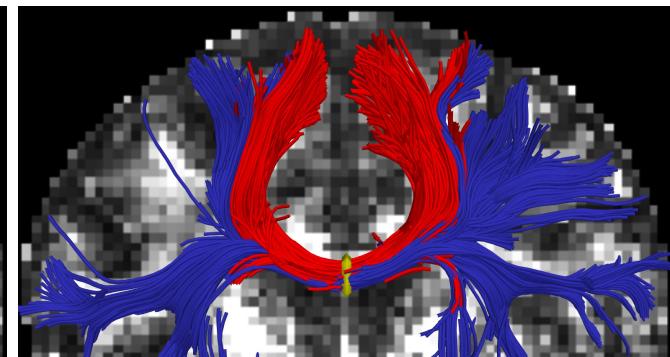
DSI [Hagmann 2005]



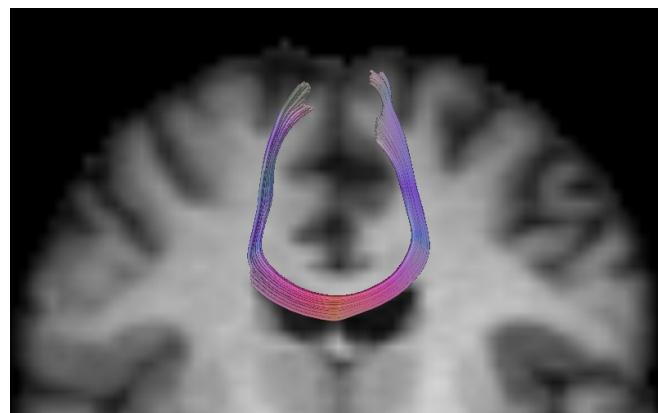
single tensor streamline



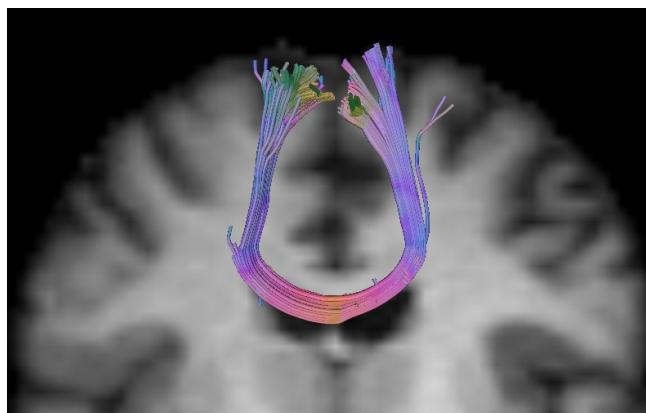
spherical harmonics



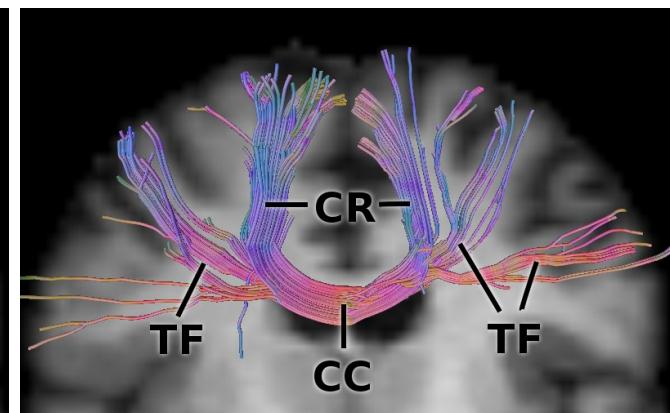
filtered two-tensor



single tensor streamline



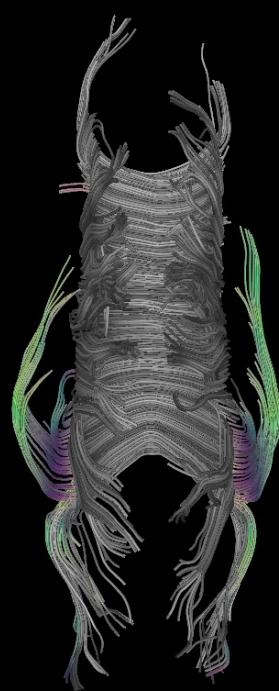
peak detection  
(our “spherical harmonics”)



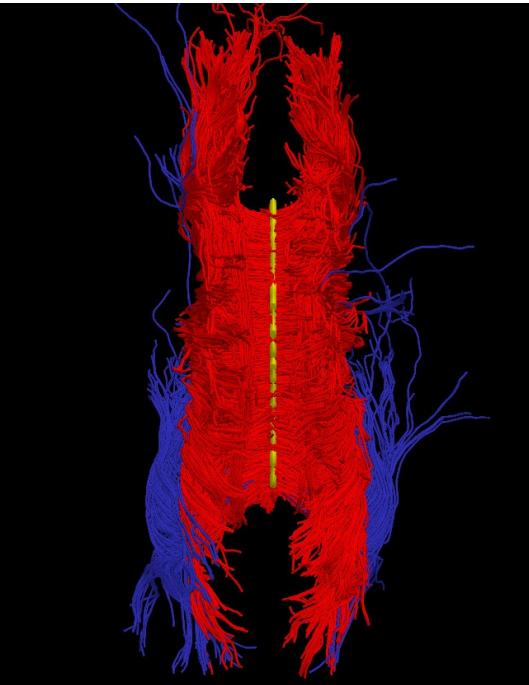
tensor decomposition

[Schultz 2008]

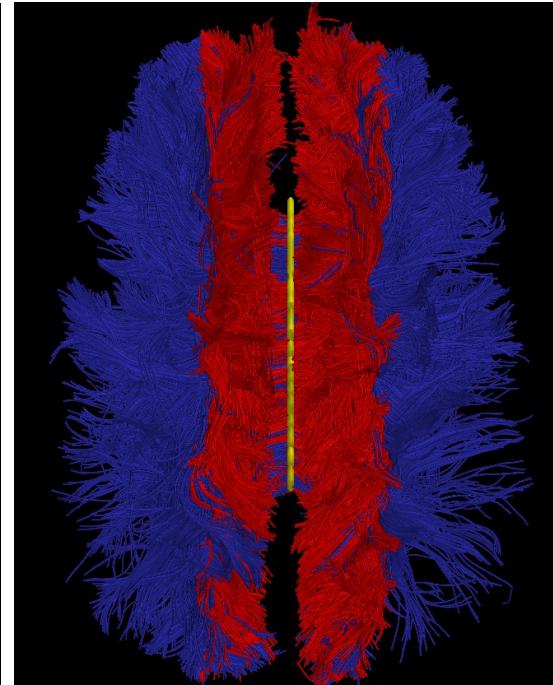
filtered two-tensor



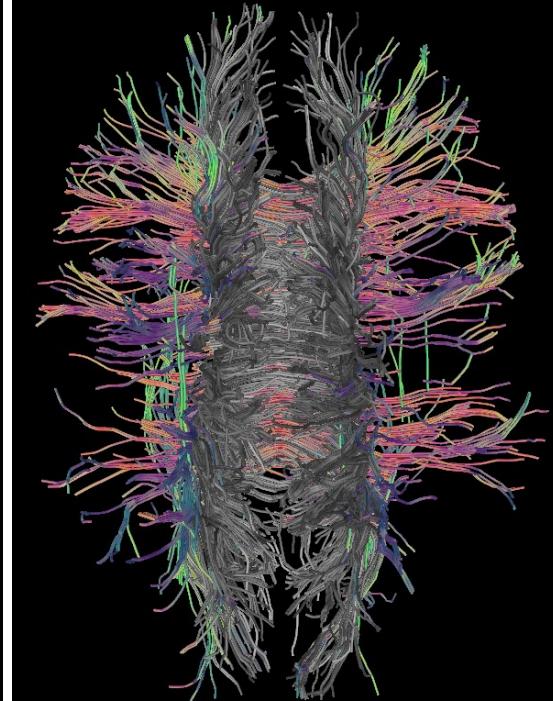
single tensor

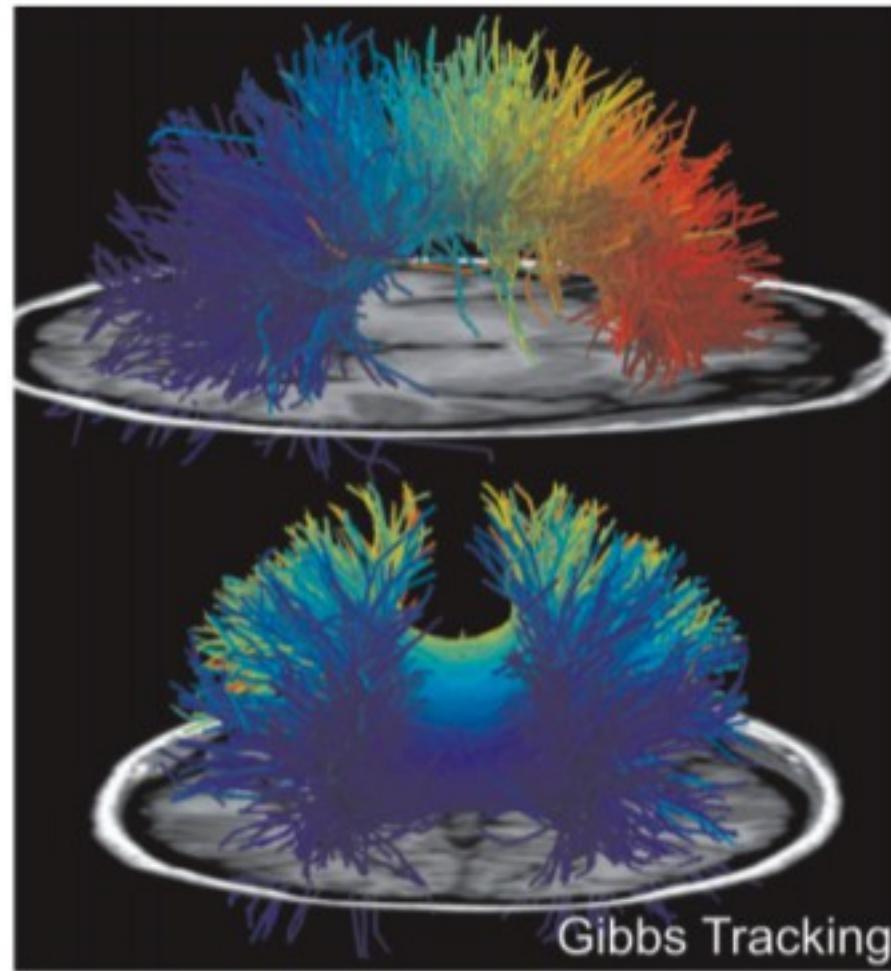


spherical harmonics



tensor decomposition  
[Schultz 2008]





[Kreher 08]

~1 month to compute

# conclusion

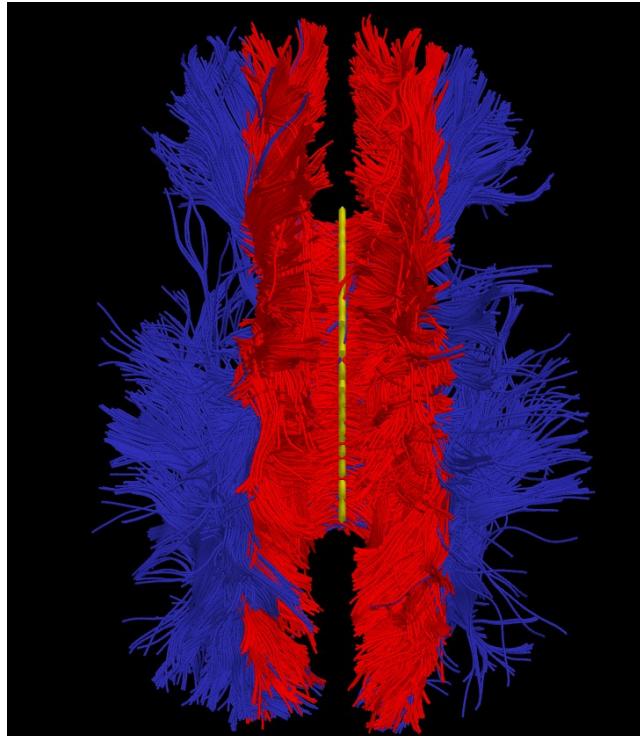
inherent coherence along the fiber

we should exploit it in the estimation

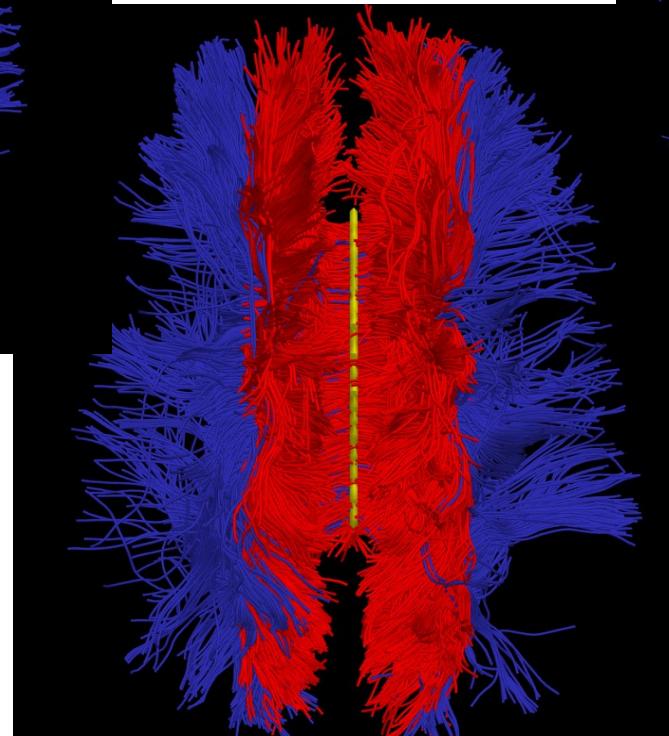
# from the proposal...

- tensors: IPMI 2009, TMI 2010
- weighted mixtures: MICCAI 2009
- validation (phantom): DMFC (MICCAI) 2010
- Slicer3 module: Slicer 3.6 (?)
- population study: DMFC (MICCAI) 2010
- local voxel model, global path model: MICCAI 2010

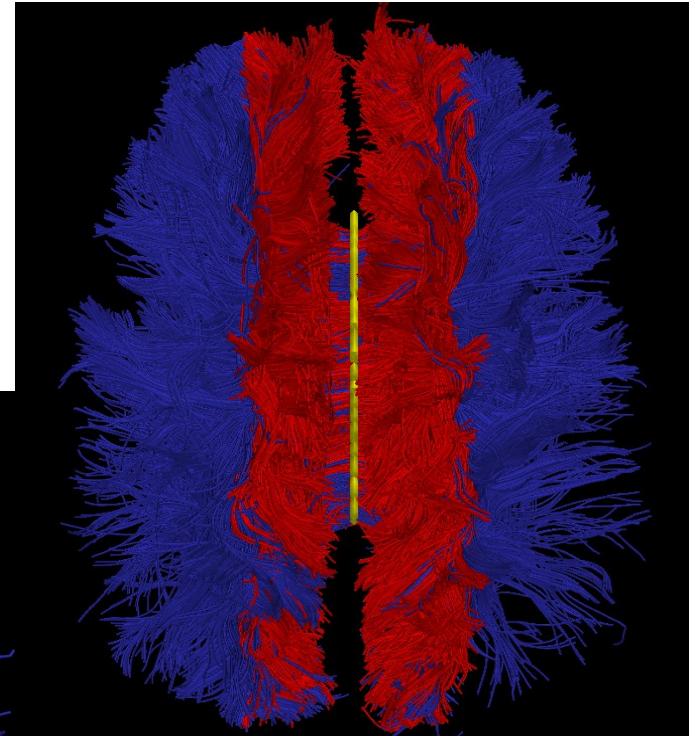
# variations



Watson functions



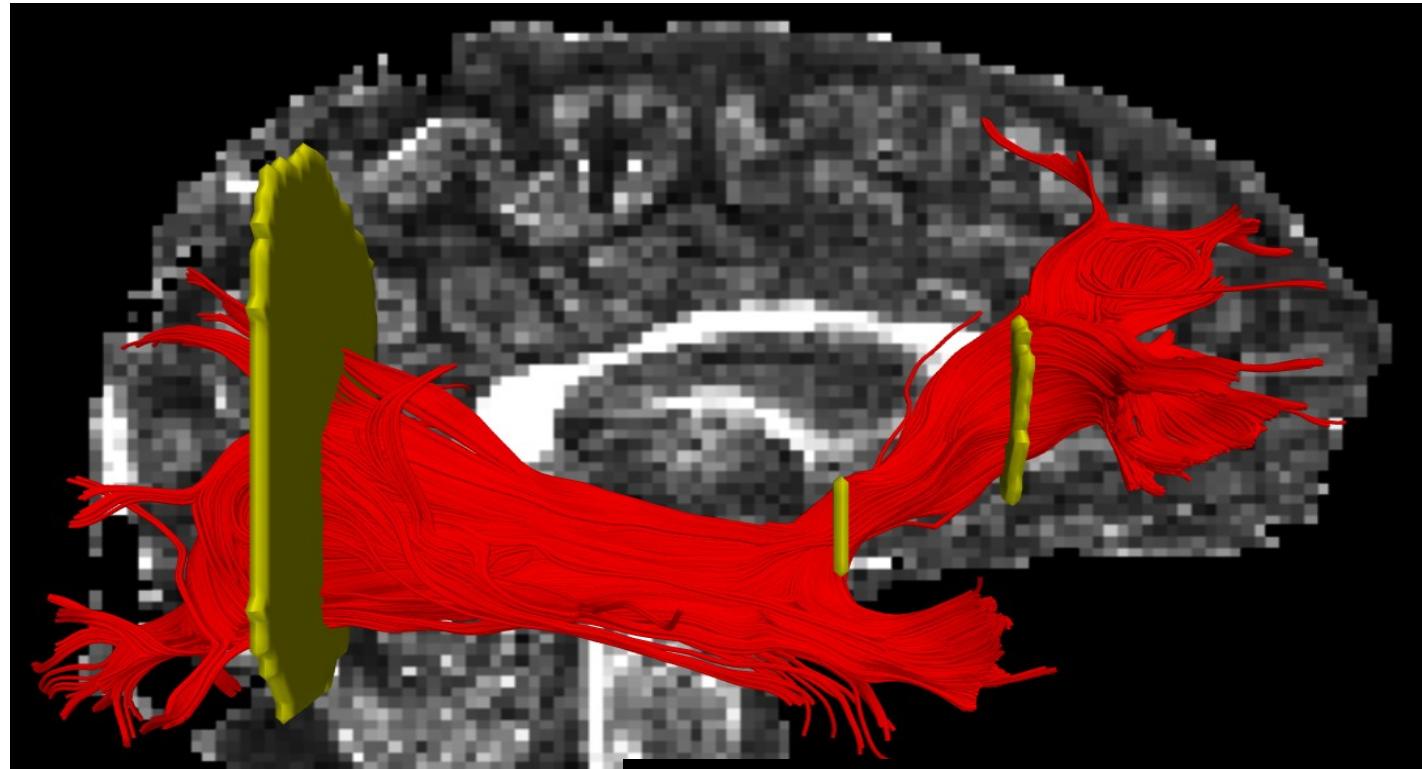
weighted models



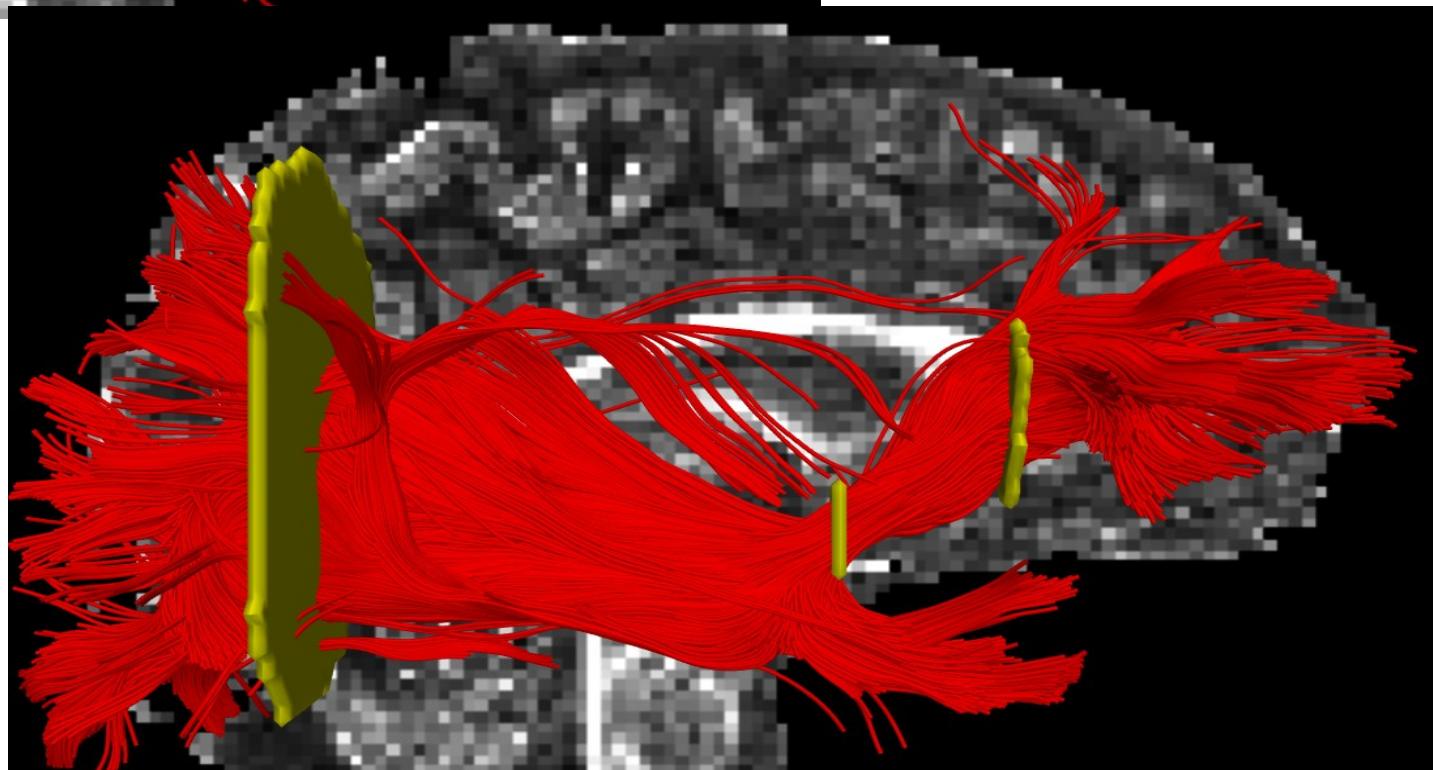
tensors

parametric:  
mixture models  
non-parametric:  
spherical harmonics  
higher-order  
tensors

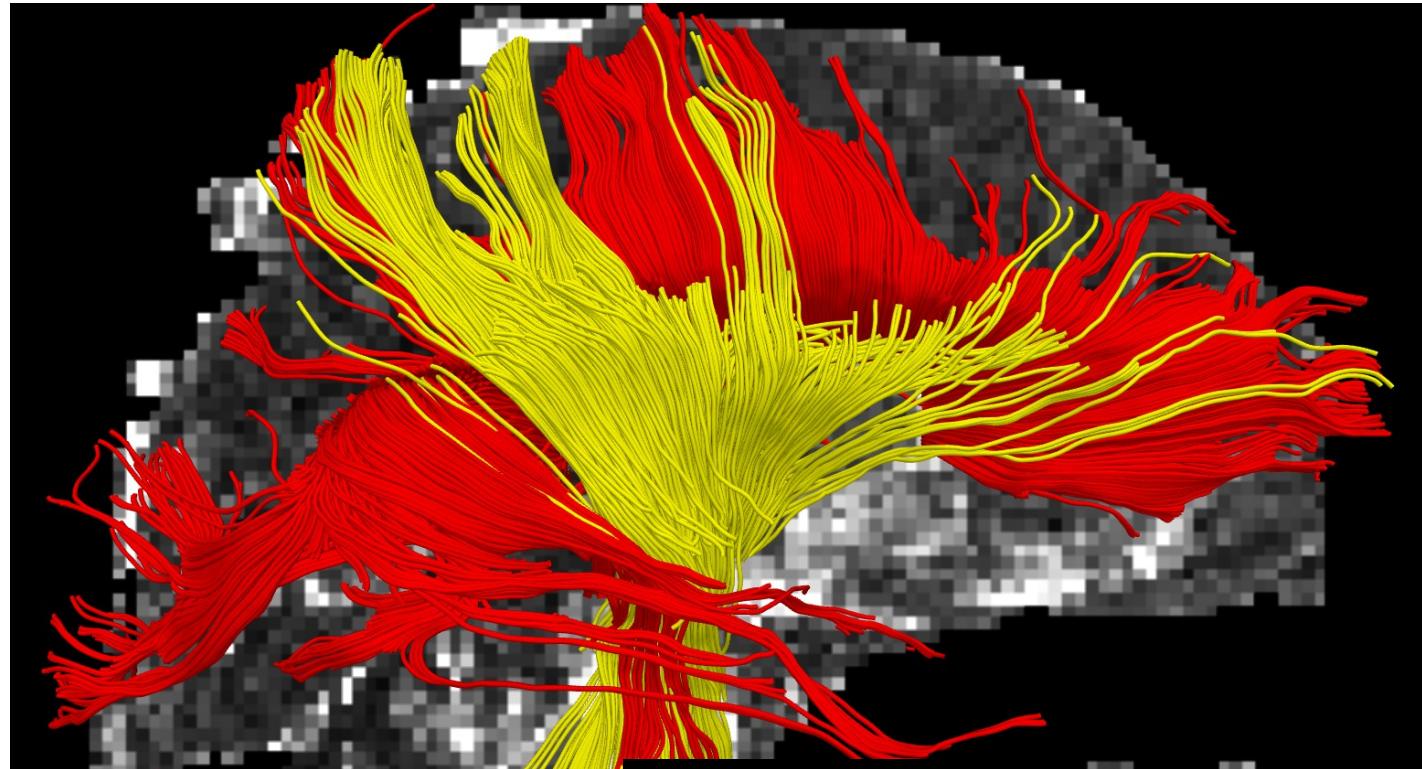
Malcolm, Shenton, Rathi. Two-tensor tractography using a constrained filter. In MICCAI, p.894–902, 2009.  
Rathi, Malcolm, Michailovich, Westin, Shenton, Bouix. Tensor-kernels for simultaneous fiber model estimation and tractography. *Magnetic Resonance in Medicine*, 64(1):138–148, 2010.



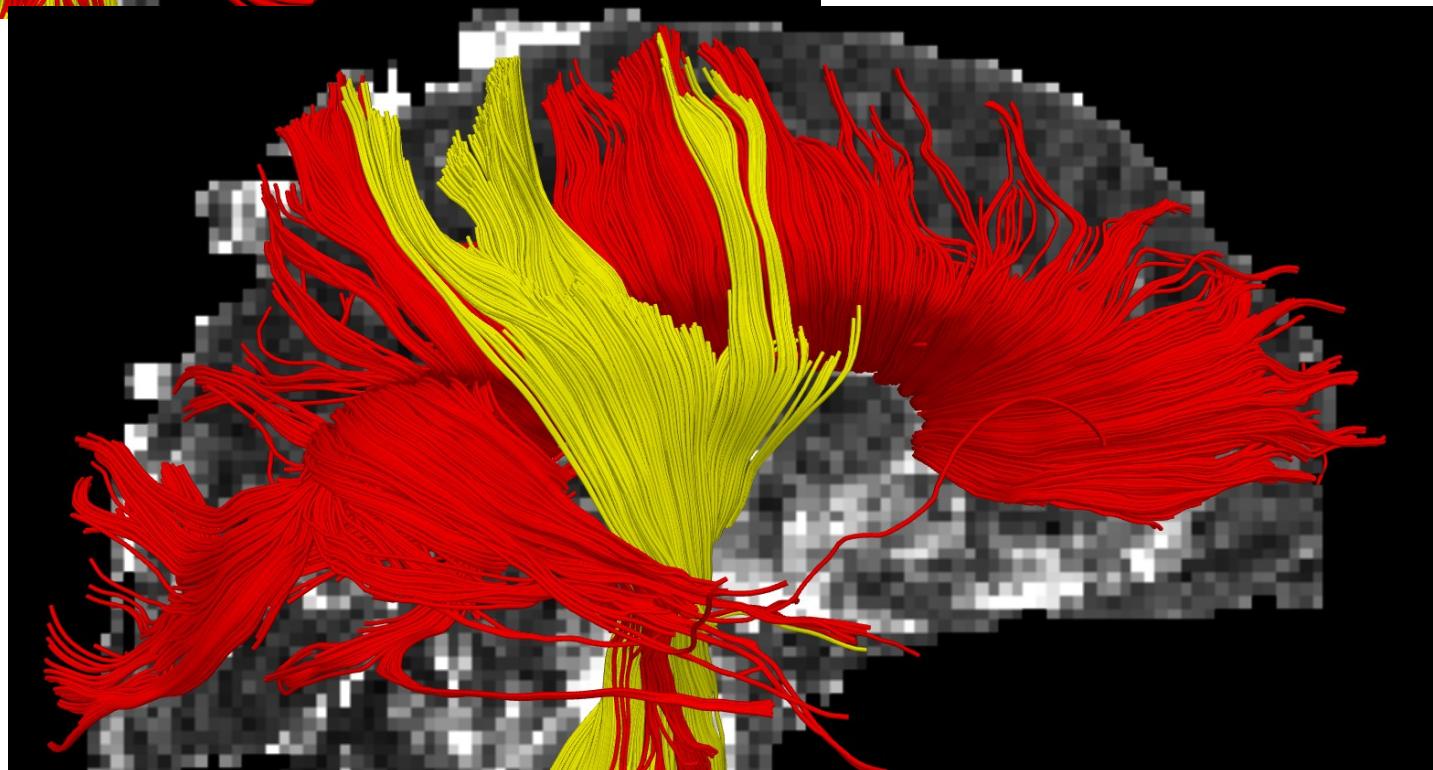
streamline  
single-tensor



filtered  
single-tensor



streamline  
single-tensor



filtered  
single-tensor

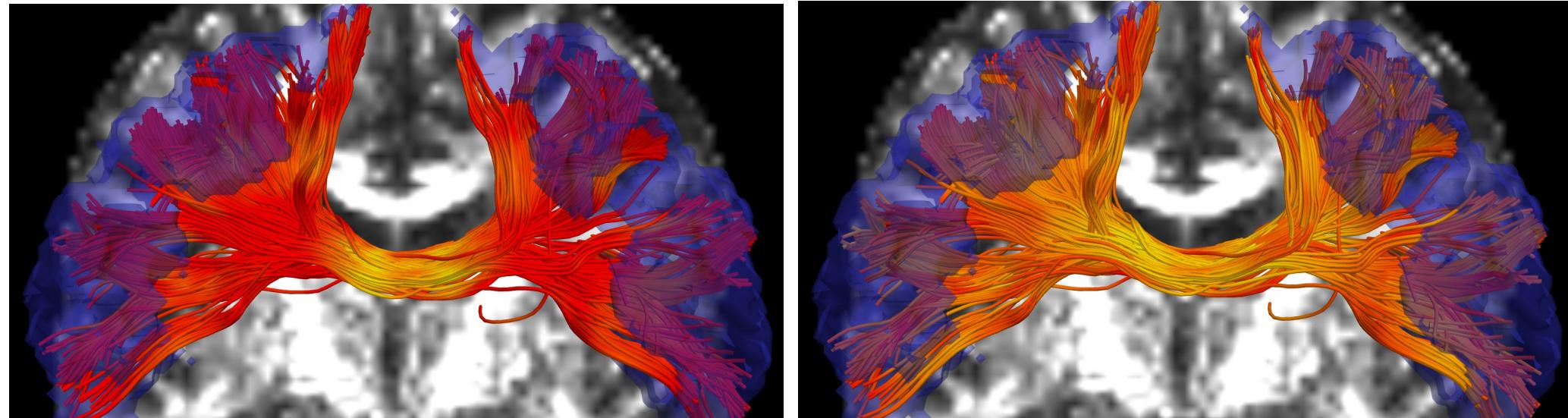
# model components one or two?

*a population study*

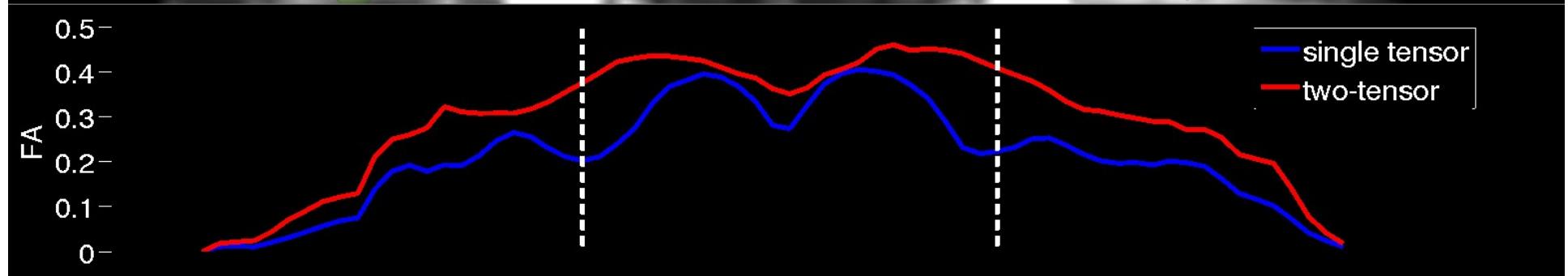
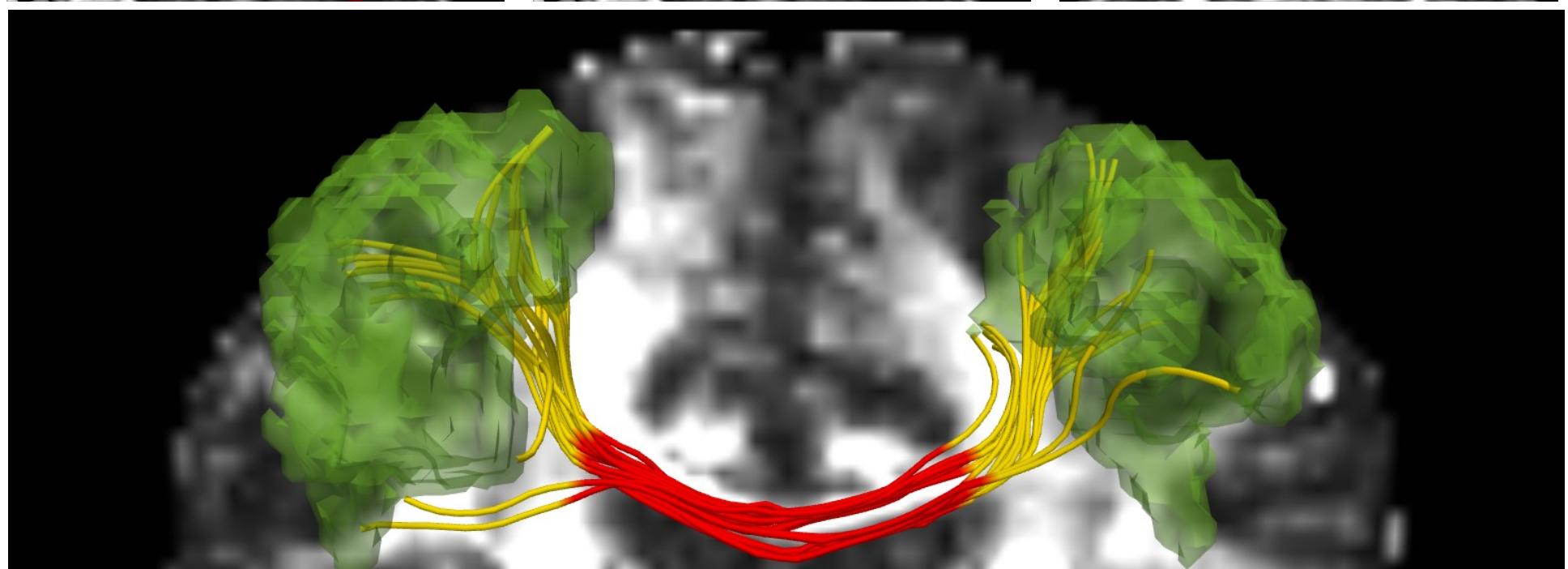
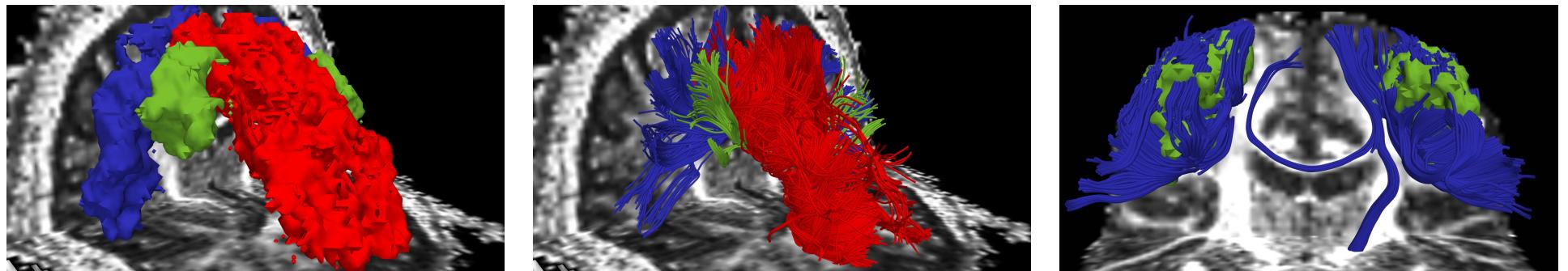
Malcolm, Kubicki, Shenton, Rathi. The effect of local fiber model on population studies. In Diffusion Modeling and Fiber Cup (in MICCAI), pages 33–40, 2009.

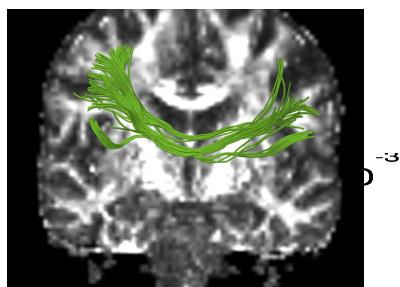
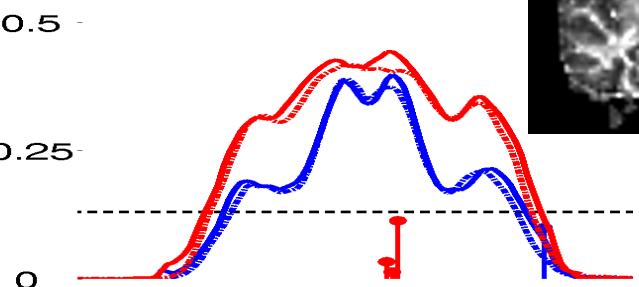
Rathi, Malcolm, Michailovich, Goldstein, Seidman, McCarley, Westin, Shenton. Biomarkers for identifying first-episode schizophrenia patients using diffusion weighted imaging. In MICCAI, volume 6361, pages 657–665, 2010.

*Experiment:* look at FA superimposed on two-tensor fibers

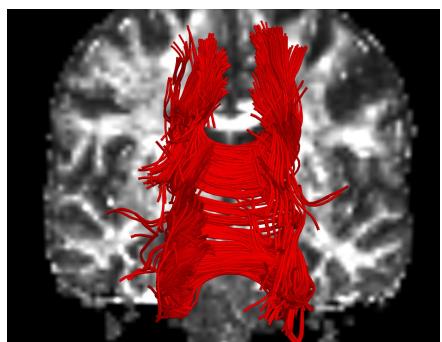
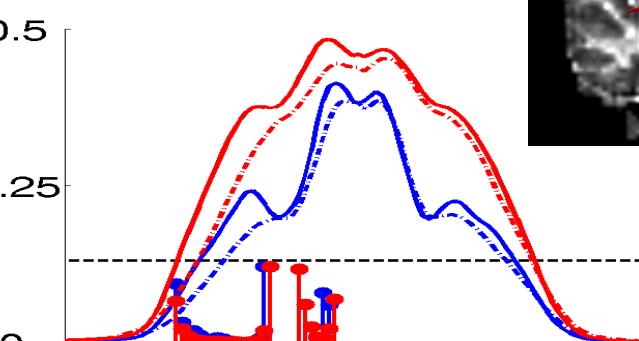
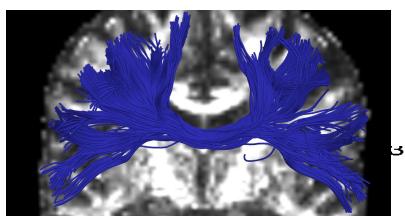
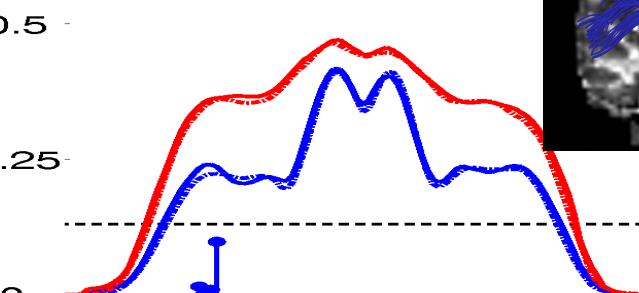
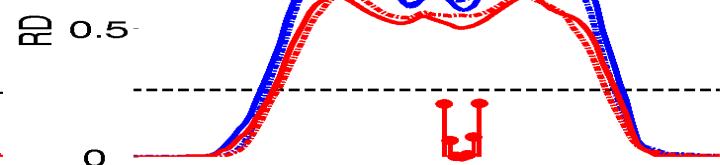


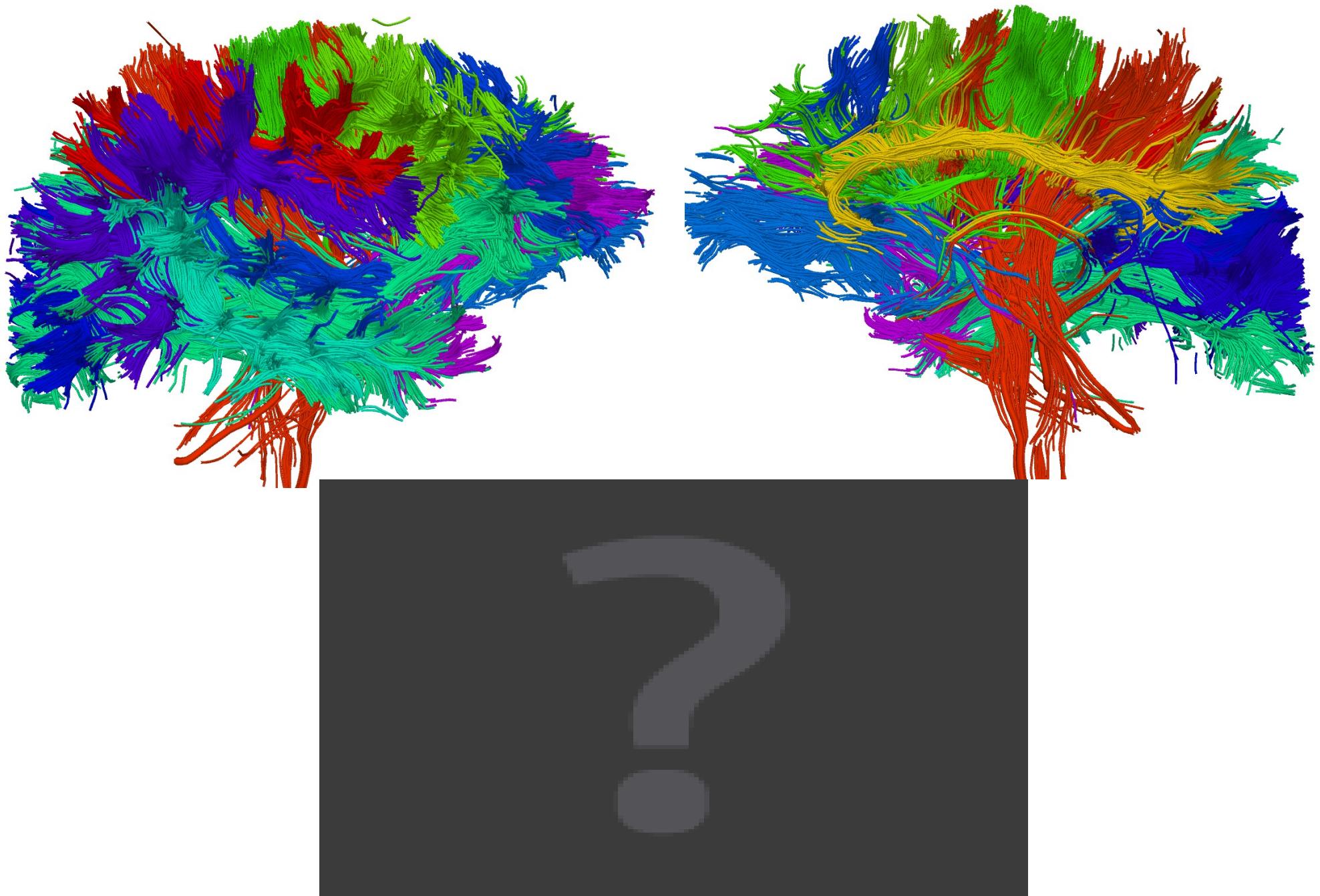
*Result:* single-tensor FA drops outside corpus callosum while two-tensor maintains higher FA on out to gray-matter





# 1T 2T





J. G. Malcolm, M. E. Shenton, and Y. Rathi. Filtered tractography: State estimation in a constrained subspace. In Diffusion Modeling and Fiber Cup (in MICCAI), pages 122–133, 2009.



end



**extra...**

# the watson approximation

J. G. Malcolm, O. Michailovich, S. Bouix, C.-F. Westin, M. E. Shenton, and Y. Rathi. A filtered approach to neural tractography using the Watson directional function. *Medical Image Analysis*, 14:58–69, 2010.

# do we need six parameters?

- for tractography, care most about direction
  - 3 parameters
- approximate diffusivity with scalar anisotropy
  - fractional anisotropy
  - 1 parameter

# Watson approximation

$$S(\mathbf{u}) = s_0 e^{-b\mathbf{u}^T D \mathbf{u}}$$

$$\begin{aligned}-b\mathbf{u}^T D \mathbf{u} &\approx -b\lambda_1 \mathbf{u}^T (\mathbf{m} \mathbf{m}^T) \mathbf{u} \\&= -b\lambda_1 (\mathbf{u}^T \mathbf{m})(\mathbf{m}^T \mathbf{u}) \\&= -k (\mathbf{u}^T \mathbf{m})^2\end{aligned}$$

$$S(\mathbf{u}) = e^{-k(\mathbf{u}^T \mathbf{m})^2}$$

principal direction     $\mathbf{m} \in \mathbb{R}^3$

scaling factor     $k \in \mathbb{R}$

4 parameters

[Rathi 09, Malcolm 10]

# model parameters

for two fibers...

...two principal directions     $m \in \mathbb{R}^3$

...two scaling factors     $k \in \mathbb{R}$

4 + 4 = 8 parameters

# model parameters

for two fibers...

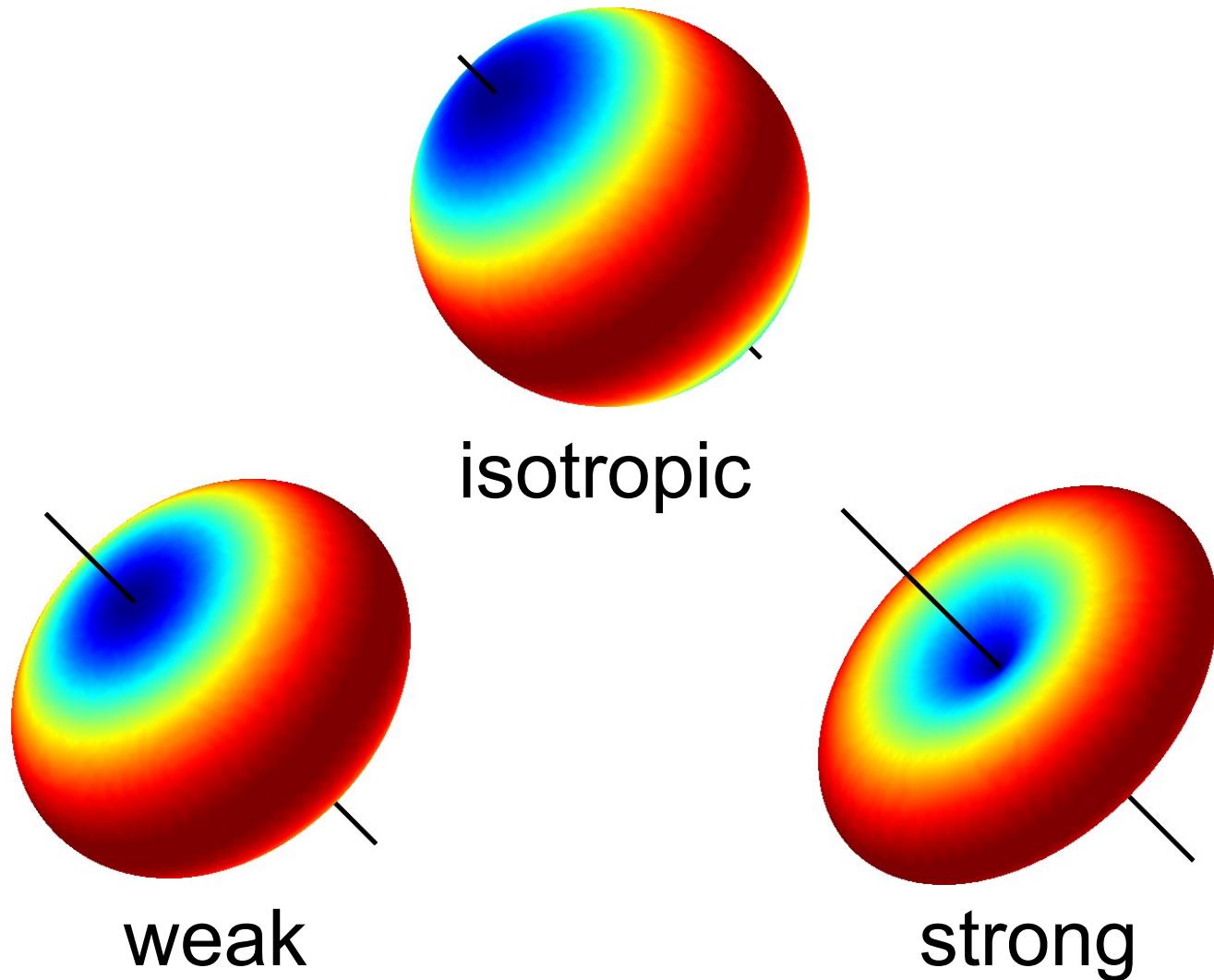
...two principal directions     $\mathbf{m} \in \mathbb{R}^3$

...two scaling factors     $k \in \mathbb{R}$

4 + 4 = 8 parameters

$$S(\mathbf{u}) = 0.5 e^{-k_1 (\mathbf{u}^T \mathbf{m}_1)^2} + 0.5 e^{-k_2 (\mathbf{u}^T \mathbf{m}_2)^2}$$

# effect of scaling factor $k$



# synthetic validation

## brute force optimization

- matching pursuit
- parametric dictionary
- noiseless signal
- discretization, noise

## spherical harmonics

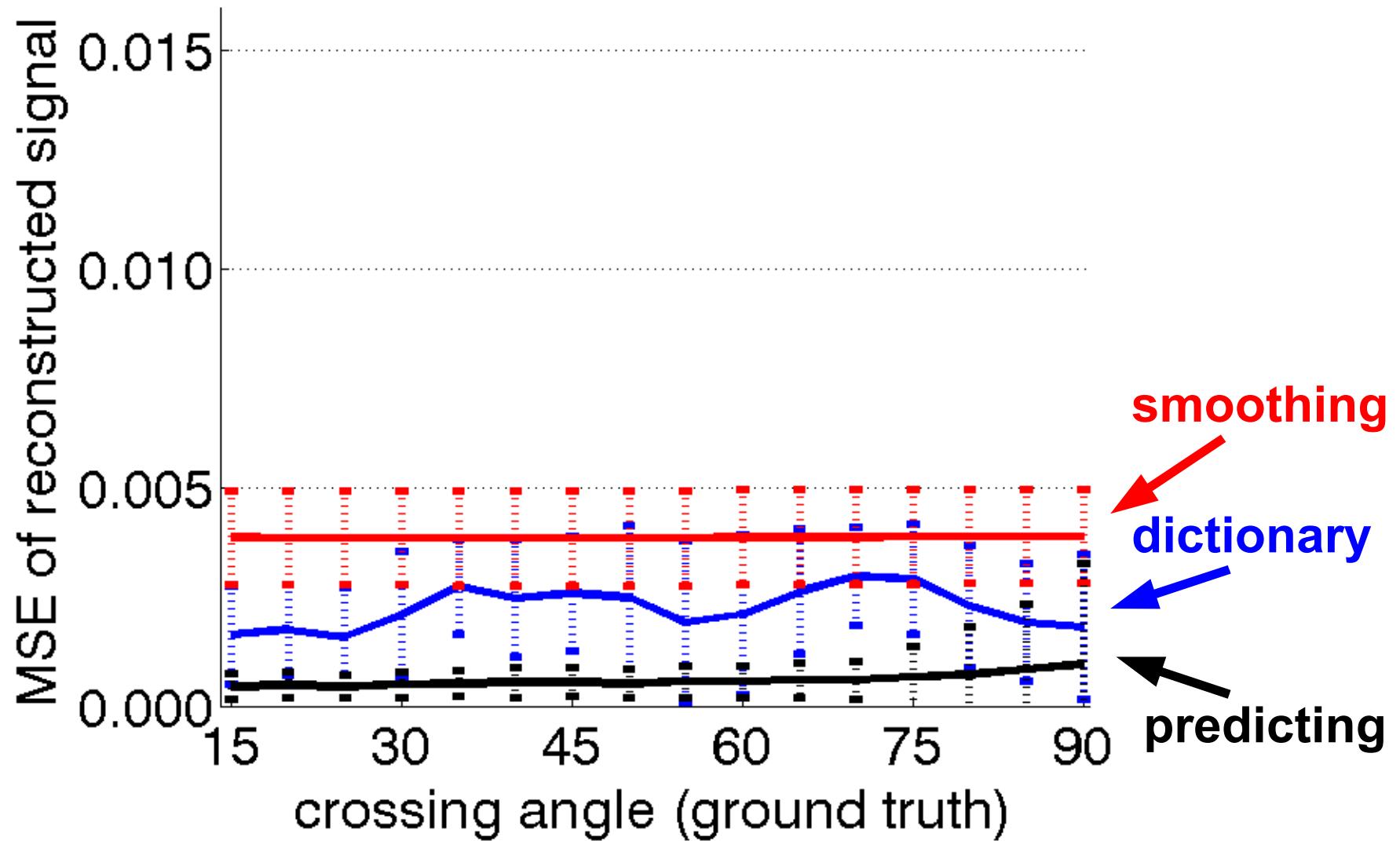
- non-parametric
- order eight (8)
- fiber sharpening for peak detection ( $L=0.006$ )

## filtered tractography

- two-fiber model
- unscented Kalman filter

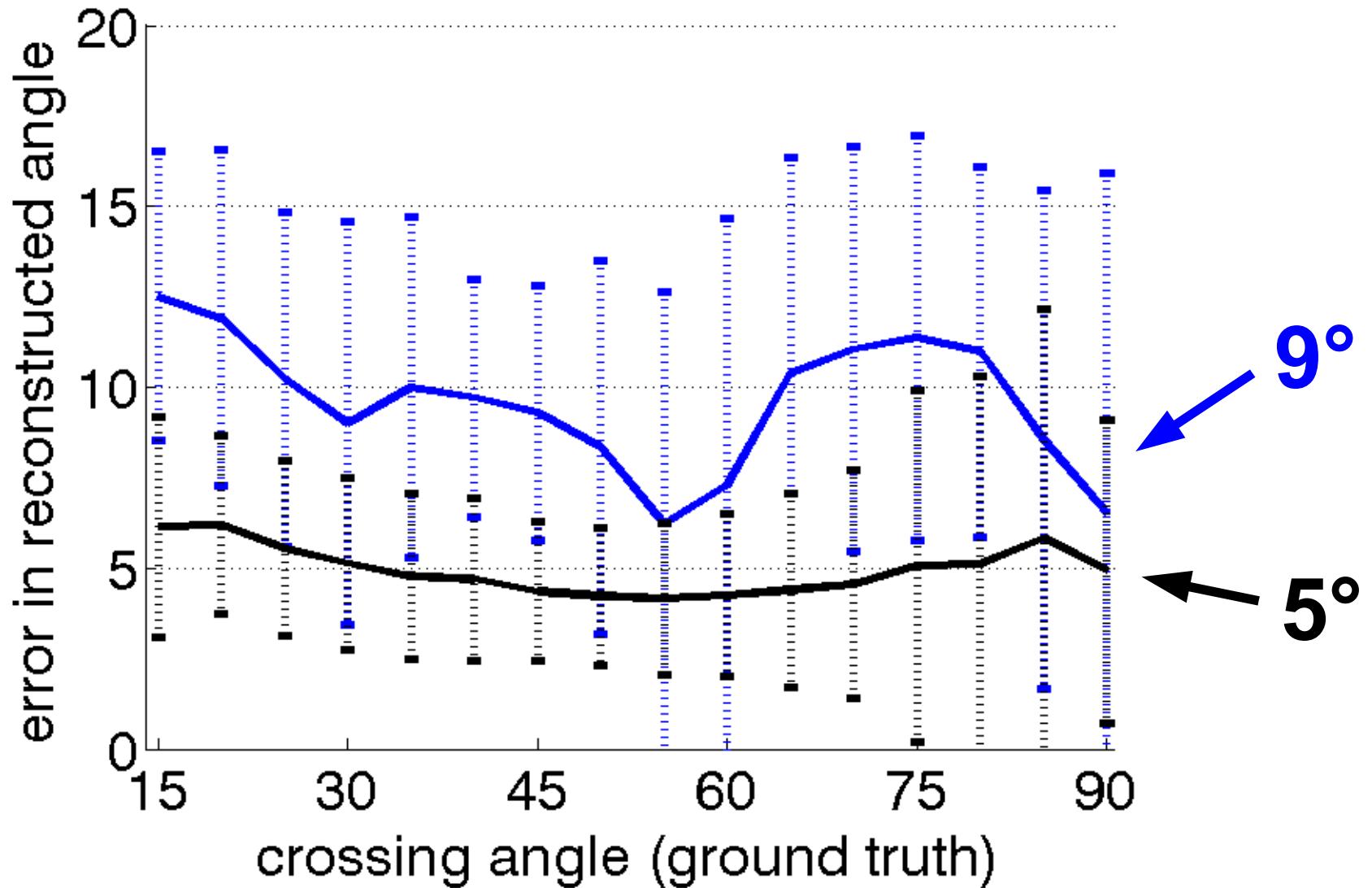
[Descoteaux 07]

# signal reconstruction error



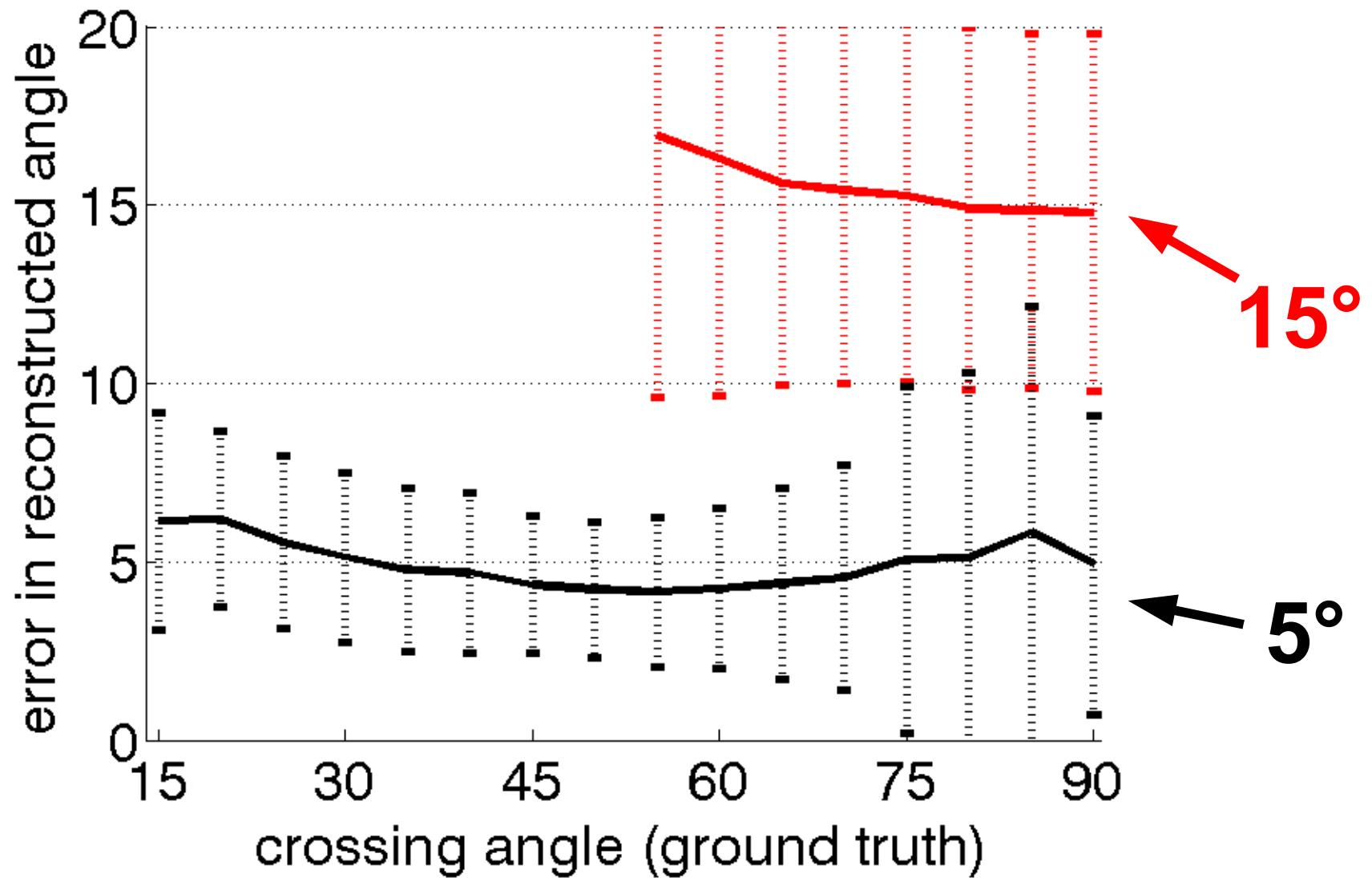
SNR  $\approx 5$ , b = 1000<sup>88</sup>

# angular reconstruction error



$\text{SNR} \approx 5, b = 1000^{89}$

# angular reconstruction error



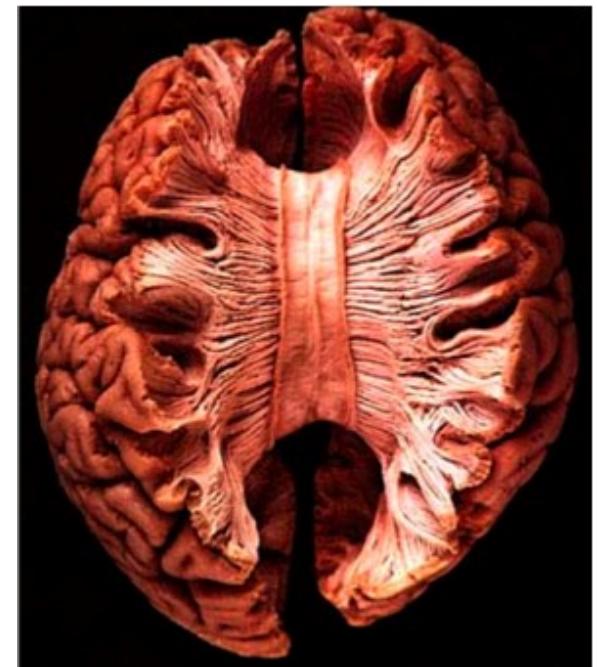
$\text{SNR} \approx 5, b = 1000^{90}$

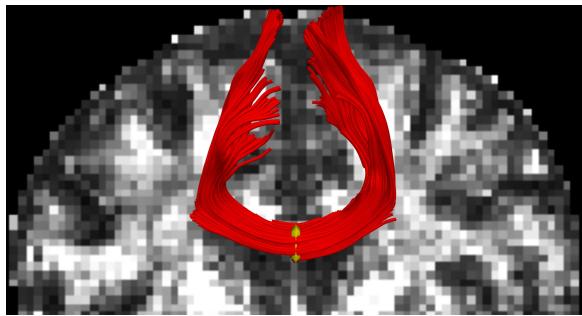


in vivo

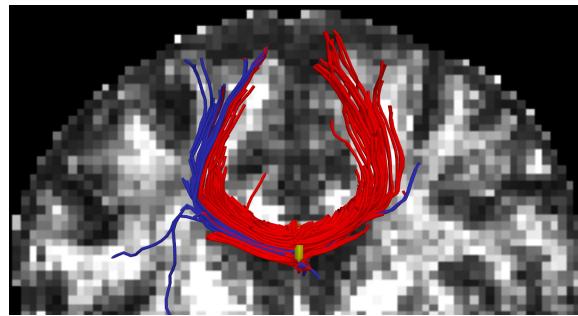


b=900  
51 directions  
1.7mm isotropic  
voxel  
17 minute scan

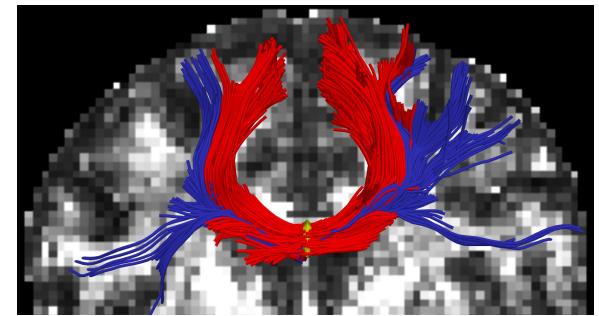




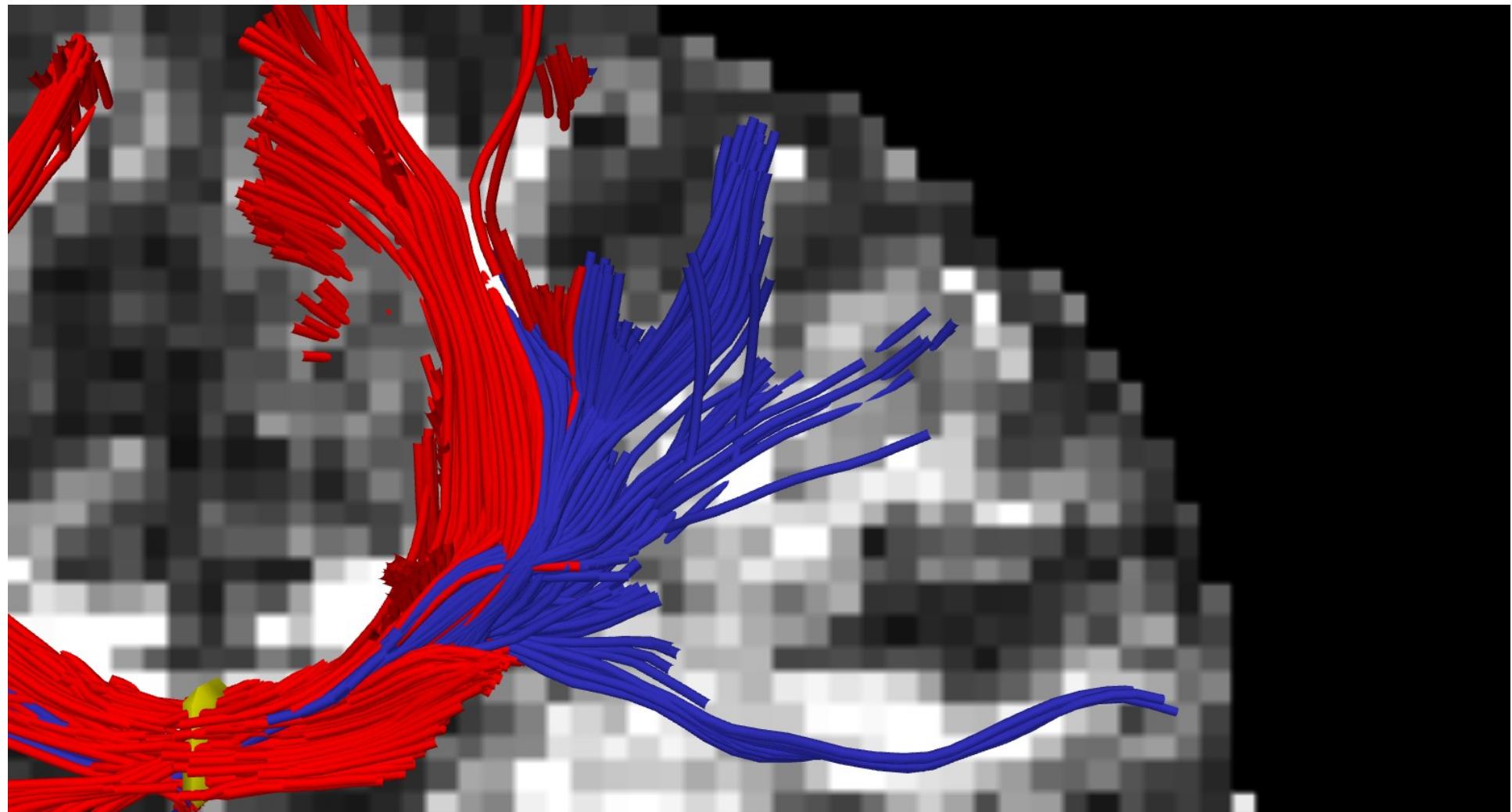
single tensor



spherical harmonics

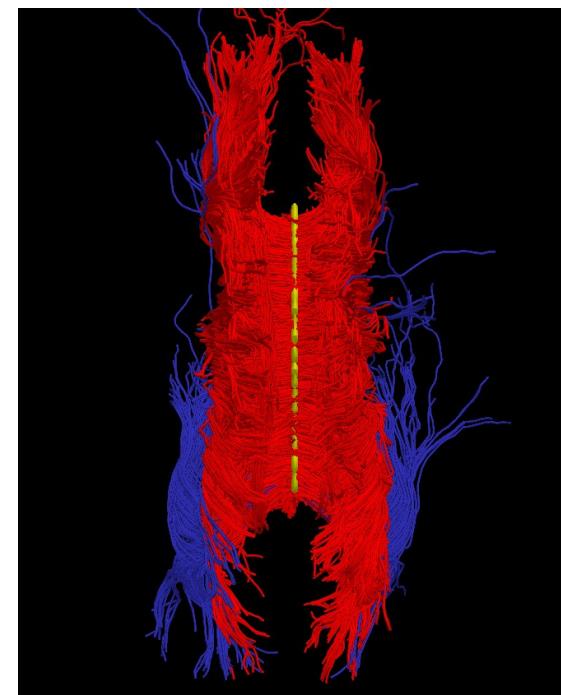
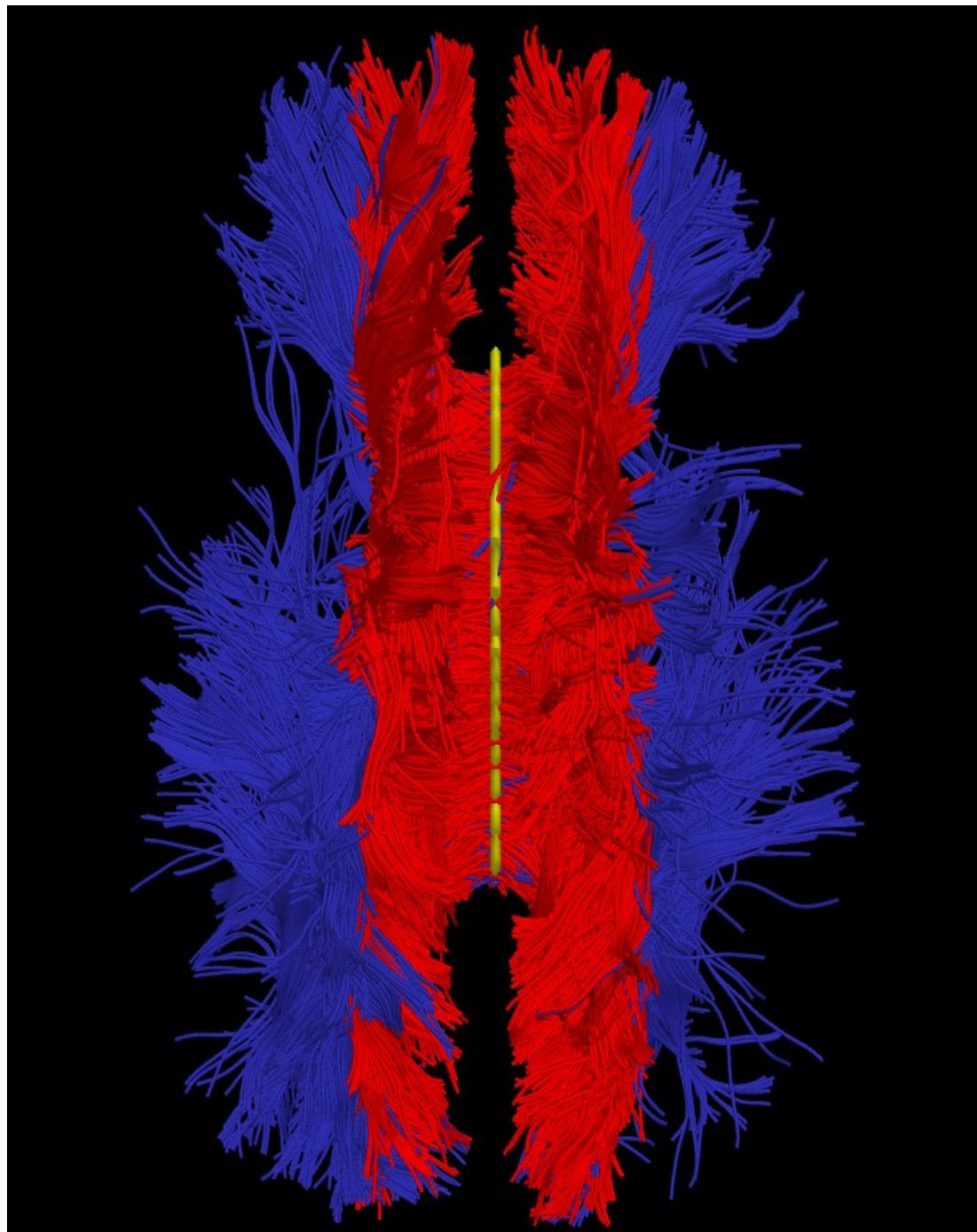


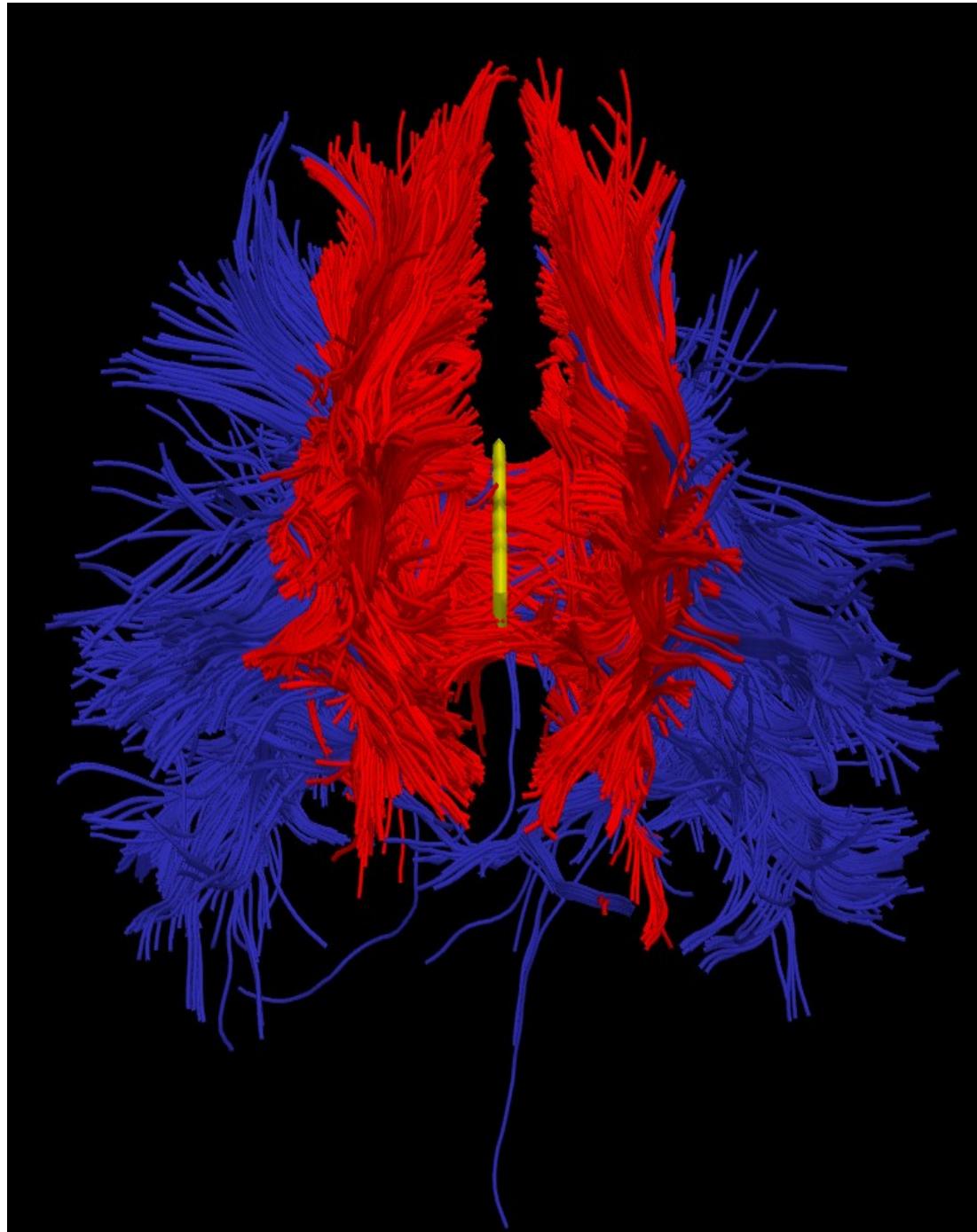
filtered two-Watson



filtered two-Watson

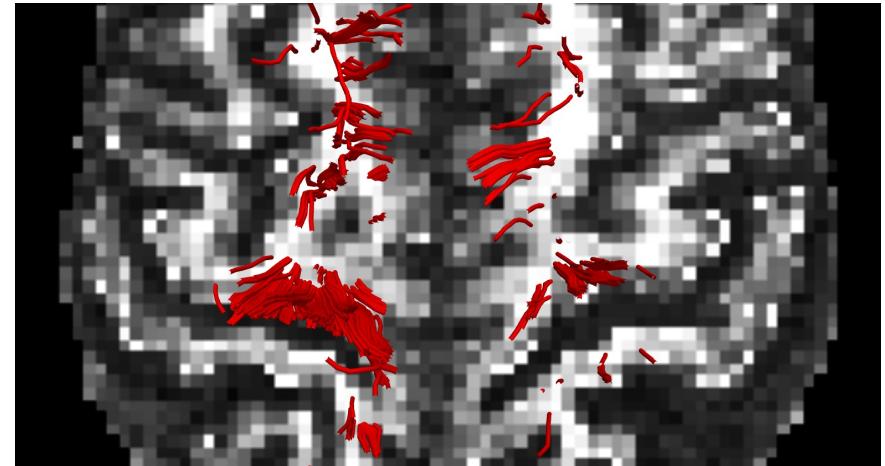
( $b = 900$ , 1.7mm, 51 directions)



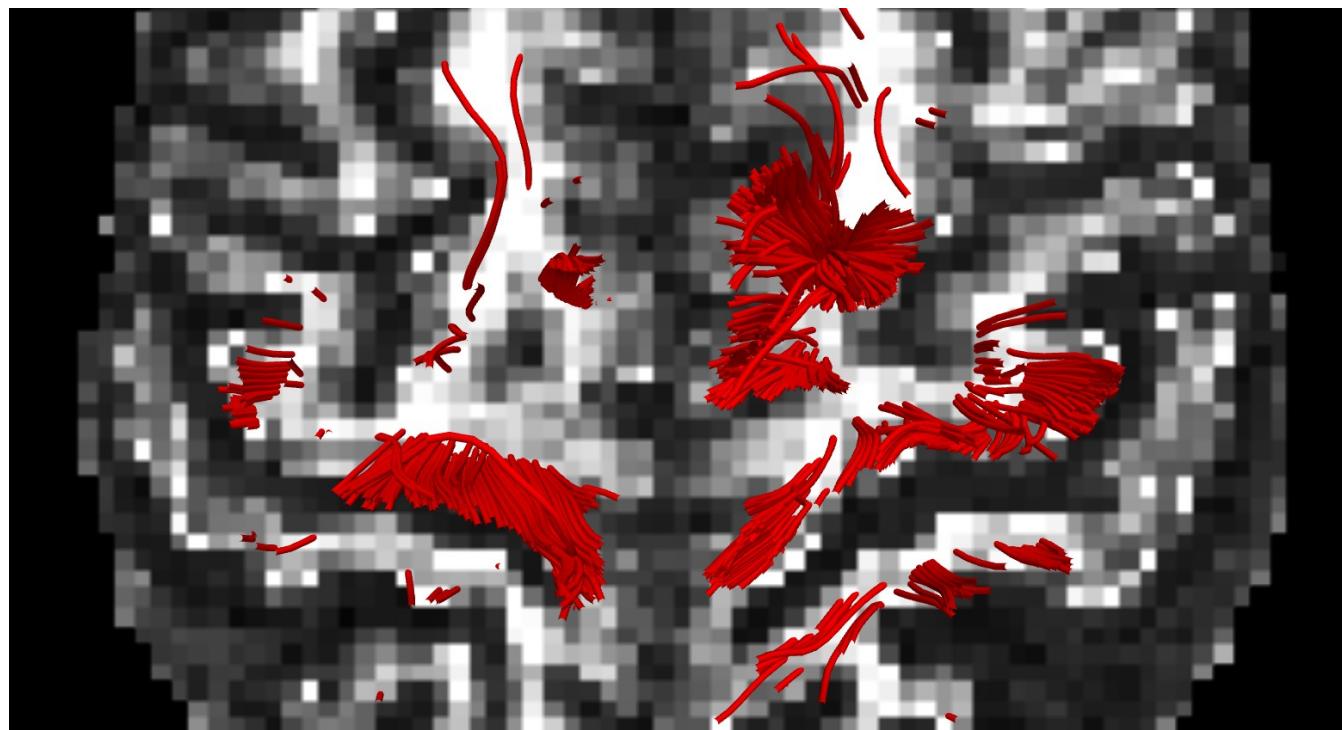




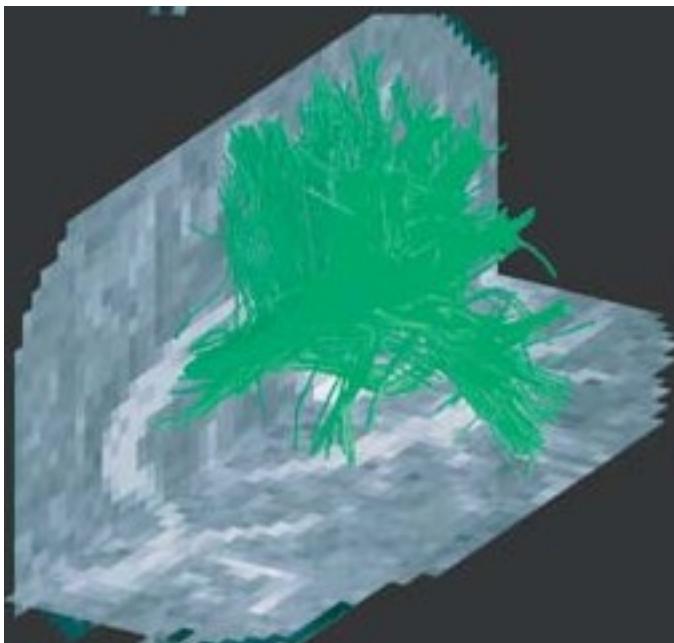
single tensor



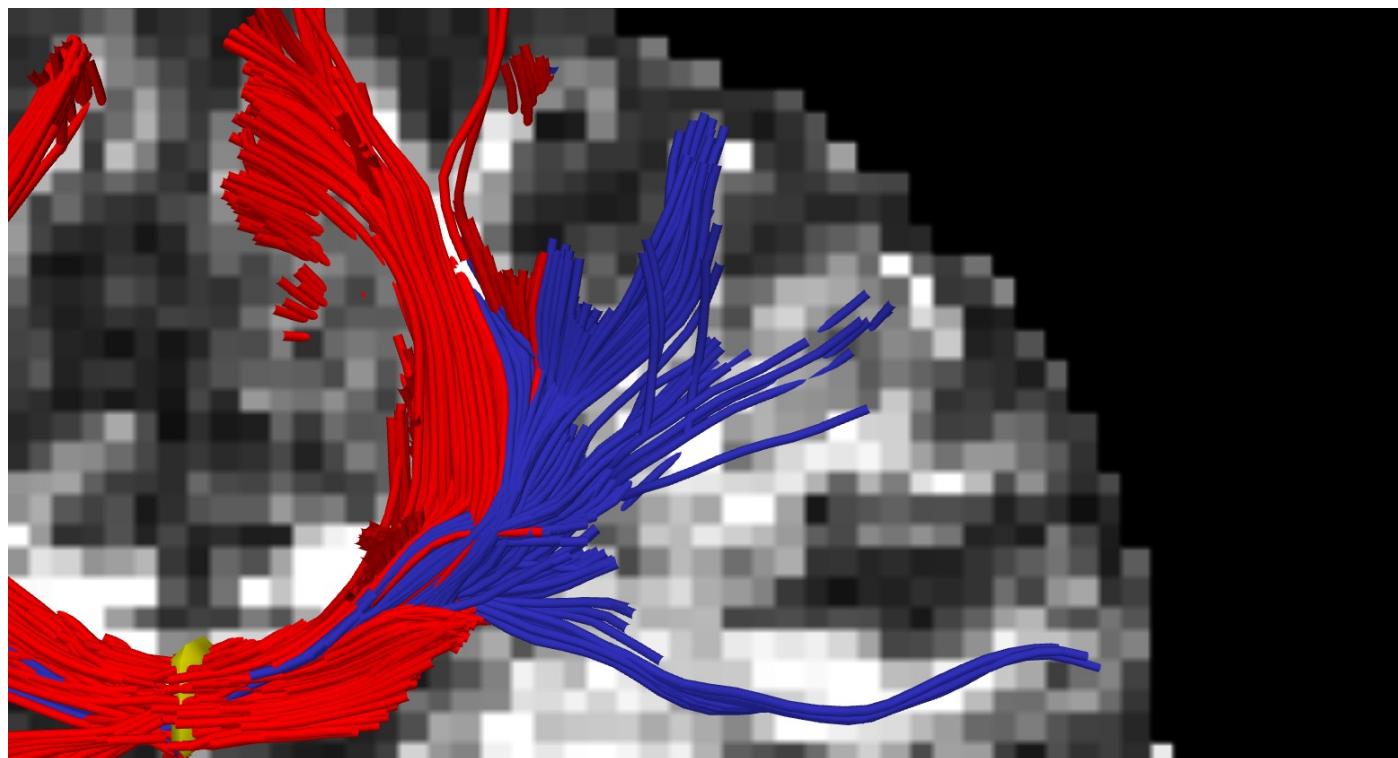
spherical harmonics



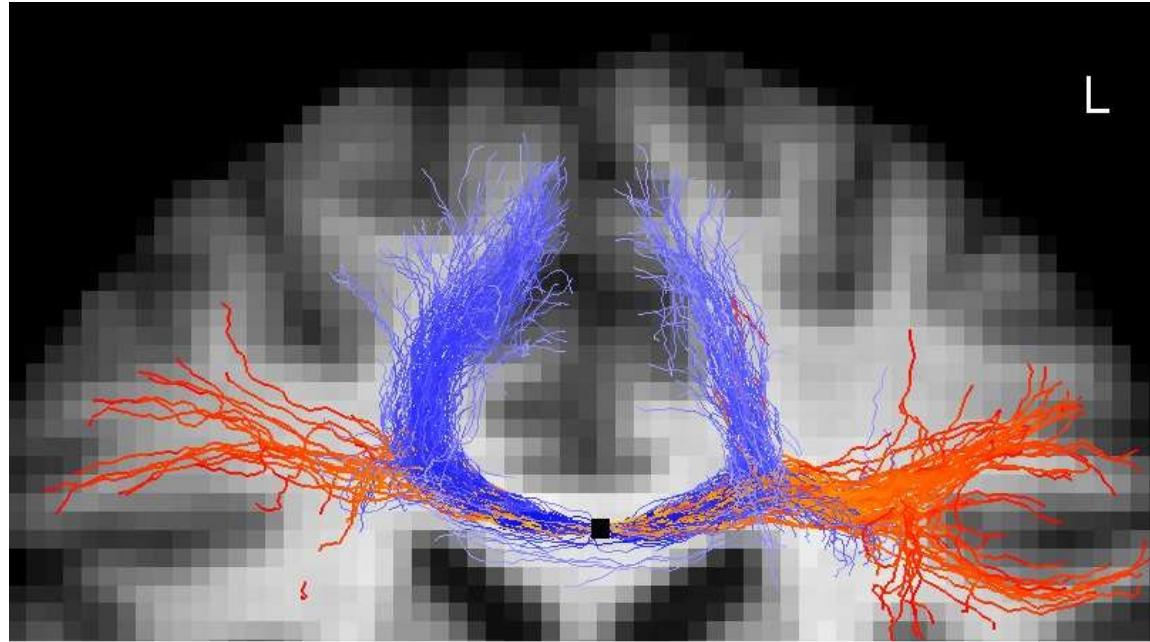
filtered two-Watson



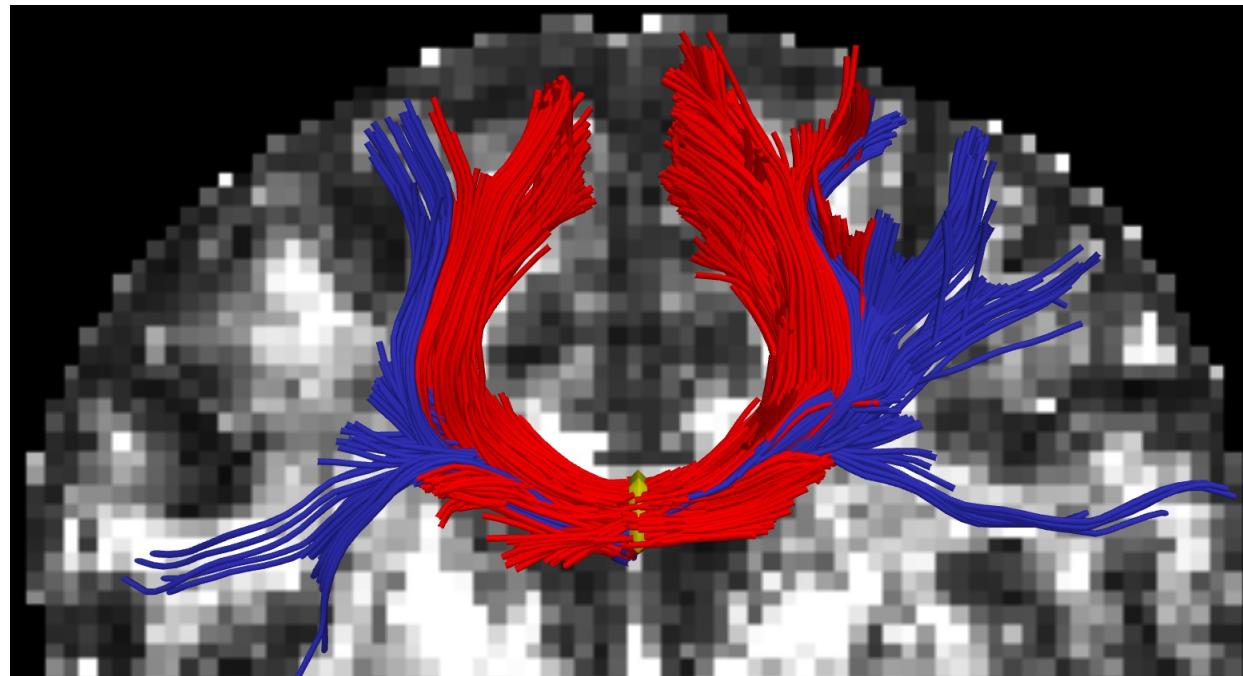
DSI [Hagmann 05]



filtered two-Watson



spherical harmonics, **probabilistic** tractography  
[Descoteaux 2009]



filtered two-Watson

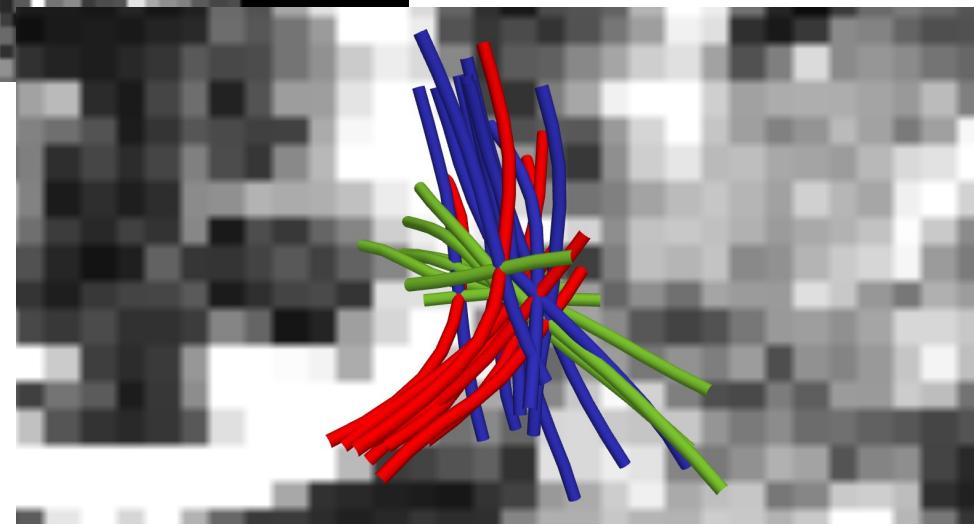
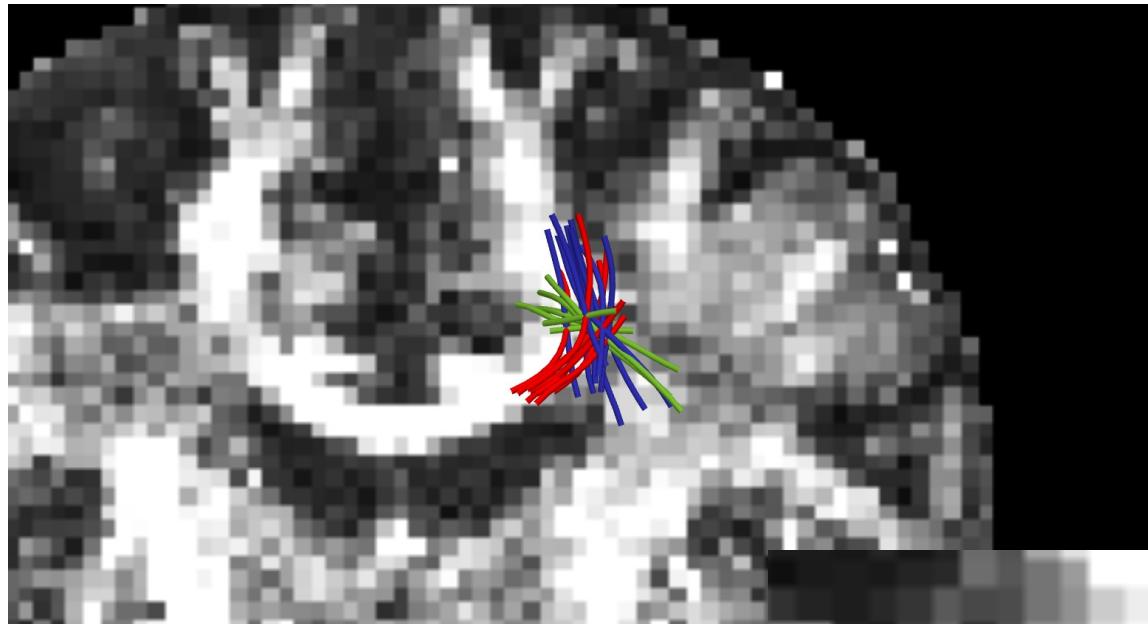
# model assumptions

*...in this proposal*

Two fibers  
single fiber: align

Fixed volume fractions  
scaling factor

model selection, non-  
parametric, ...



---

**Algorithm 1** Unscented Kalman Filter

---

- 1: Form weighted sigma points  $\mathbf{X}_t = \{w_i, \mathbf{x}_i\}_{i=0}^{2n}$  around current mean  $\mathbf{x}_t$  and covariance  $P_t$  with scaling factor  $\zeta$

$$\mathbf{x}_0 = \mathbf{x}_t \quad \mathbf{x}_i = \mathbf{x}_t + [\sqrt{\zeta P_t}]_i \quad \mathbf{x}_{i+n} = \mathbf{x}_t - [\sqrt{\zeta P_t}]_i$$

- 2: Predict the new sigma points and observations

$$\mathbf{X}_{t+1|t} = f[\mathbf{X}_t] \quad \mathbf{Y}_{t+1|t} = h[\mathbf{X}_{t+1|t}]$$

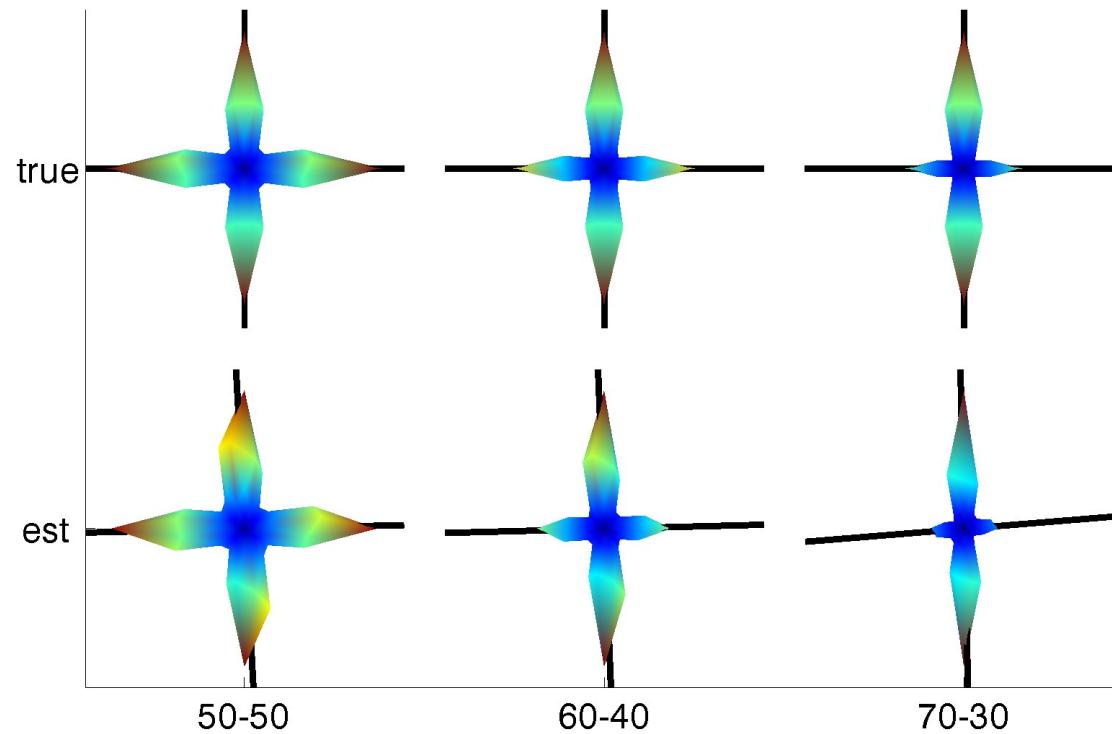
- 3: Compute weighted means and covariances, e.g.

$$\bar{\mathbf{x}}_{t+1|t} = \sum_i w_i \mathbf{x}_i \quad P_{xy} = \sum_i w_i (\mathbf{x}_i - \bar{\mathbf{x}}_{t+1|t})(\mathbf{y}_i - \bar{\mathbf{y}}_{t+1|t})^T$$

- 4: Update estimate using Kalman gain  $K$  and scanner measurement  $\mathbf{y}_t$

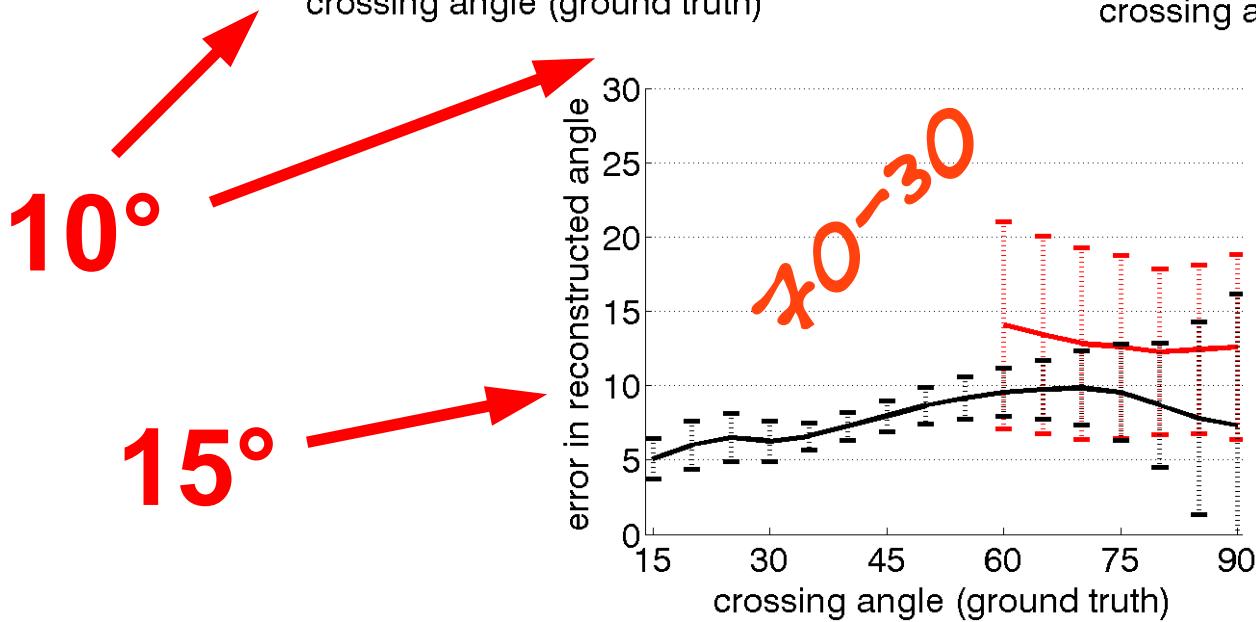
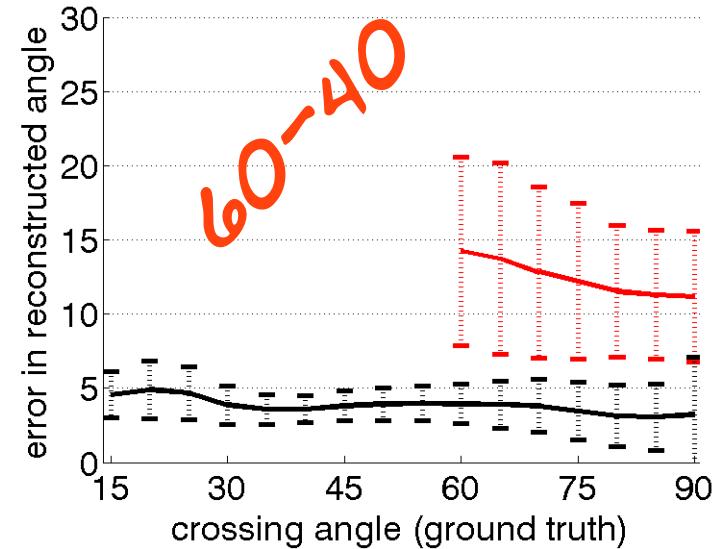
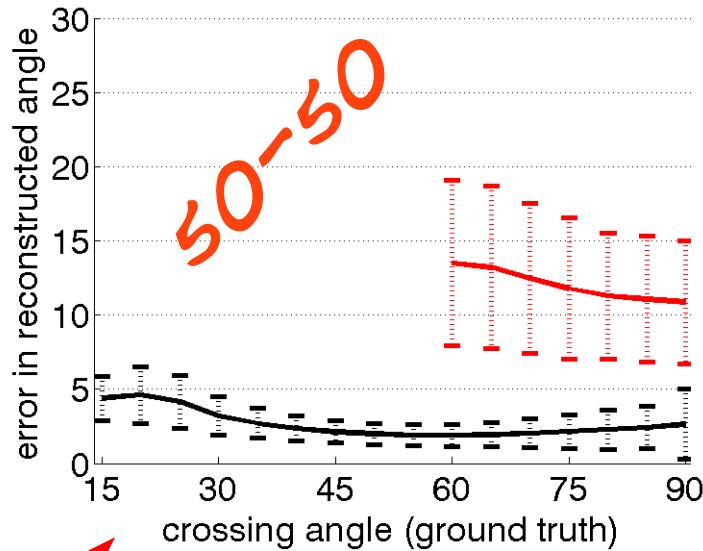
$$\mathbf{x}_{t+1} = \bar{\mathbf{x}}_{t+1|t} + K(\mathbf{y}_t - \bar{\mathbf{y}}_{t+1|t}) \quad P_{t+1} = P_{xx} - K P_{yy} K^T \quad K = P_{xy} P_{yy}^{-1}$$

# volume fractions



$\text{SNR} \approx 10, b = 1000^{101}$

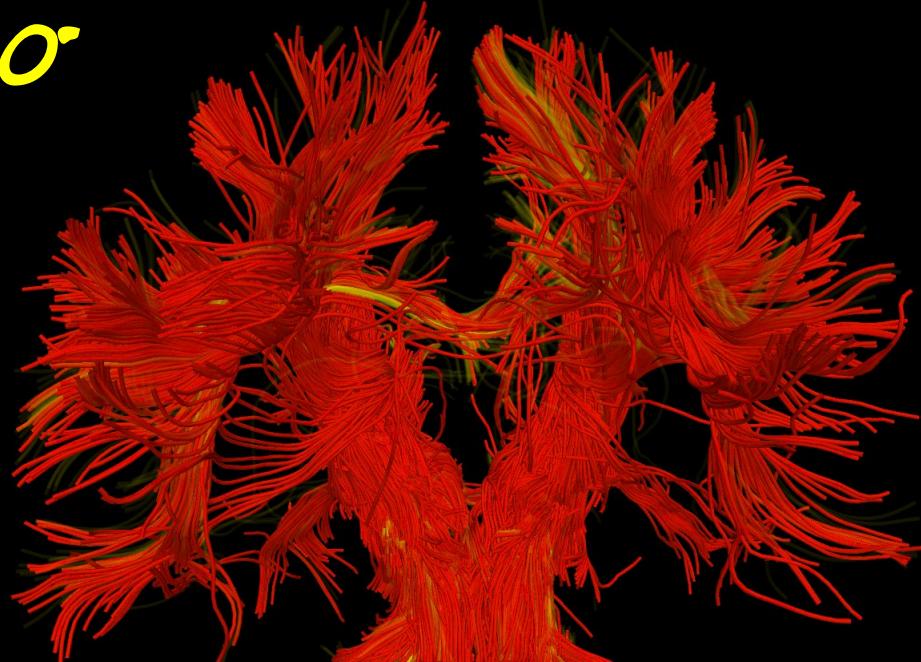
# volume fractions



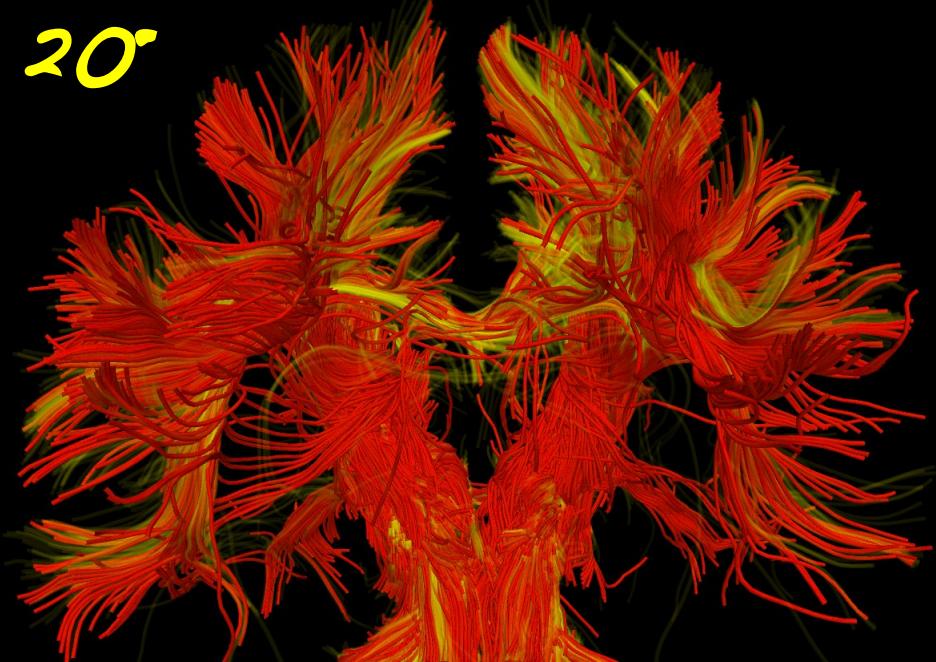
filter  
deteriorates

$\text{SNR} \approx 10, b = 1000^{102}$

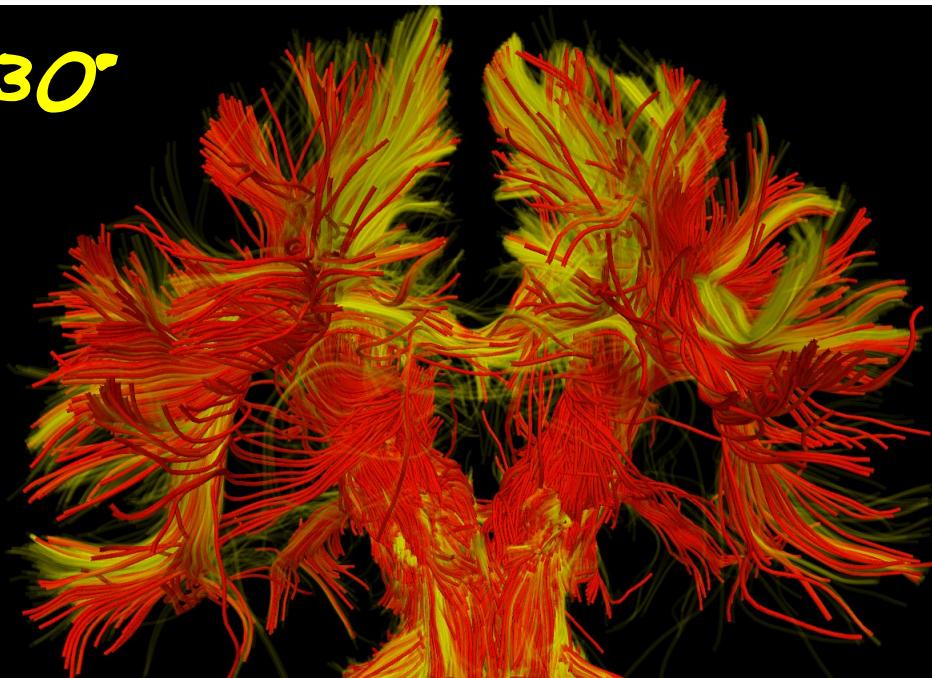
$10^\circ$



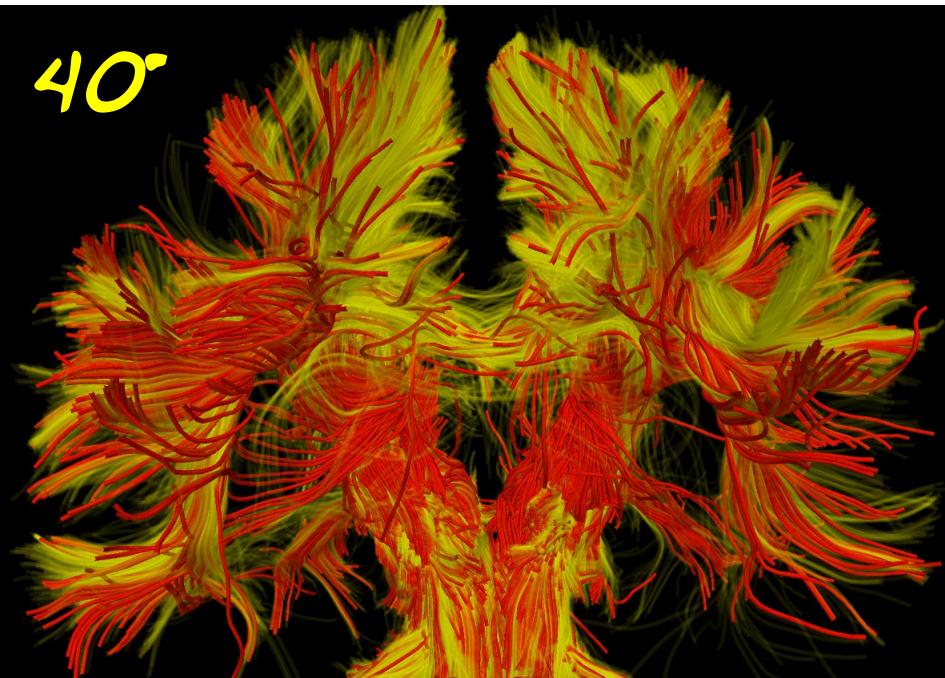
$20^\circ$



$30^\circ$

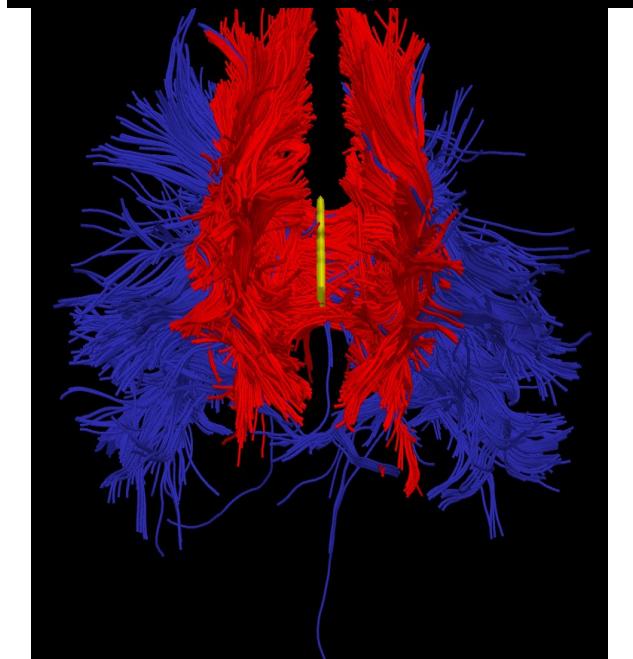
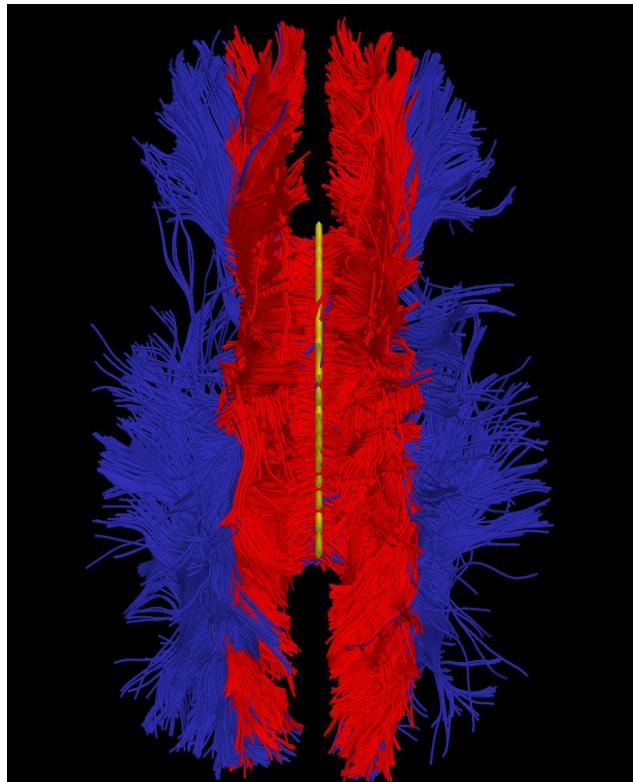


$40^\circ$



primary

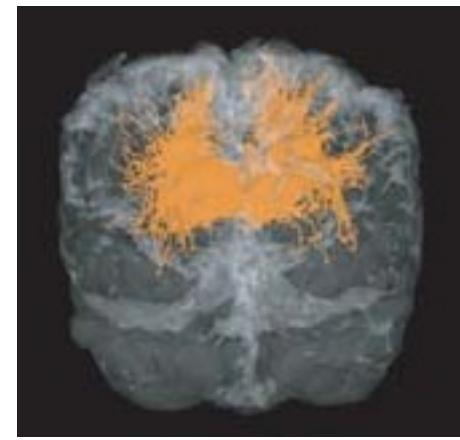
branches



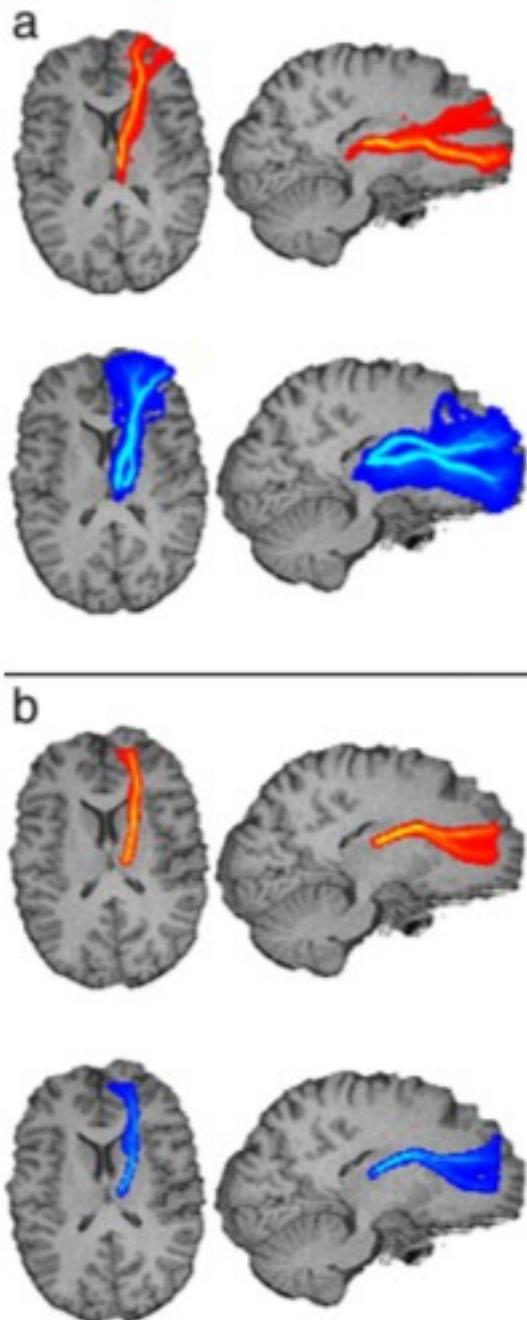
filtered two-Watson



DSI (Hagmann 2005)

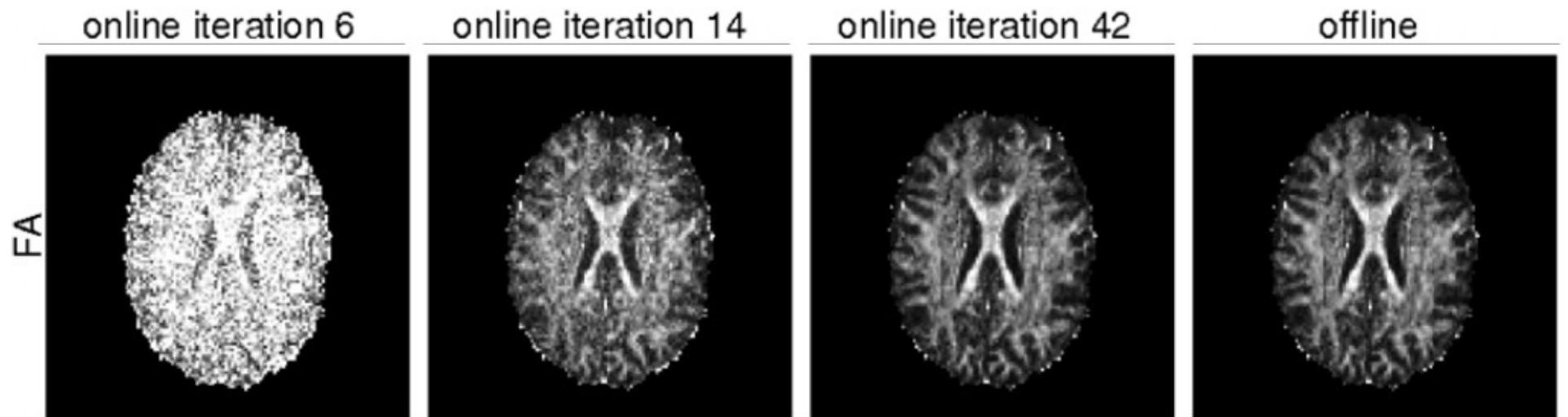


DSI (Hagmann 2005)

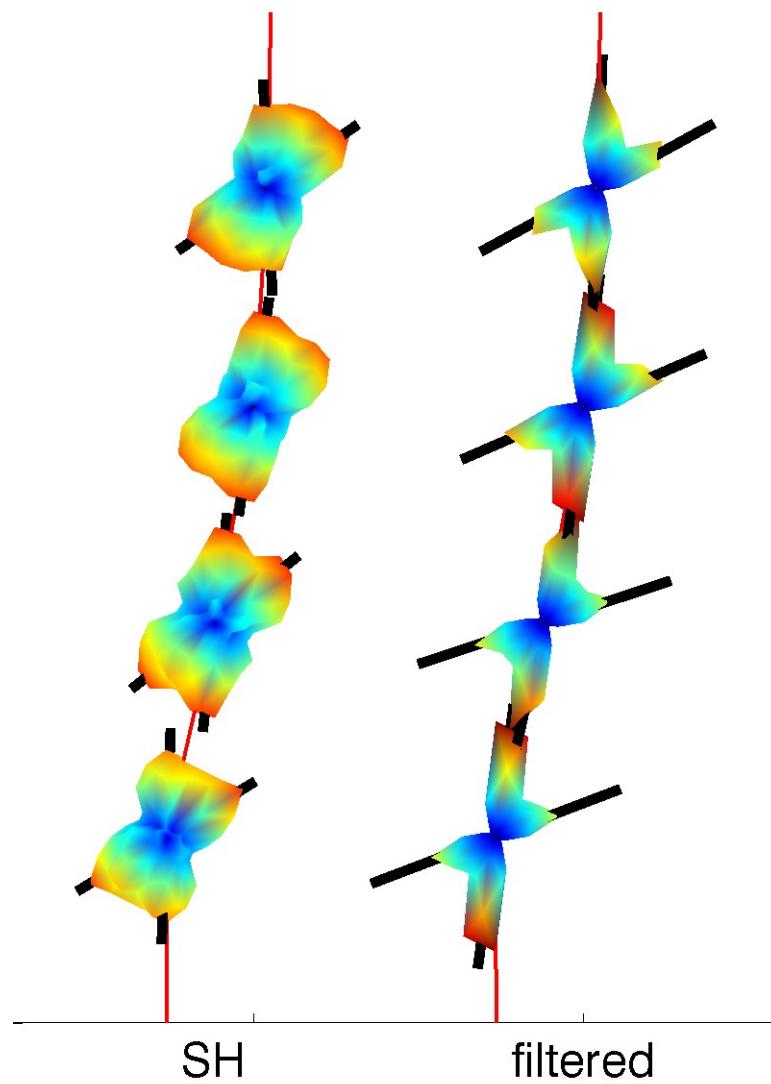
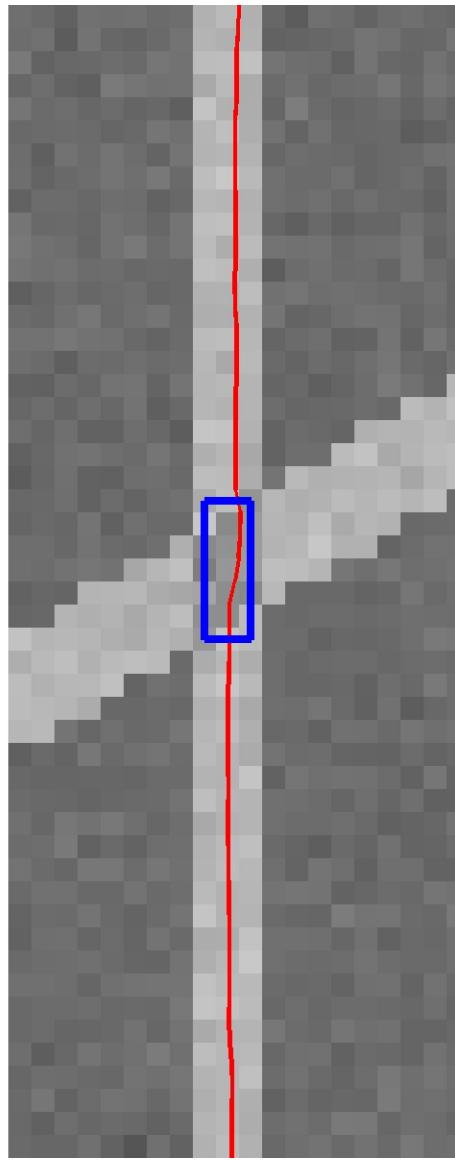


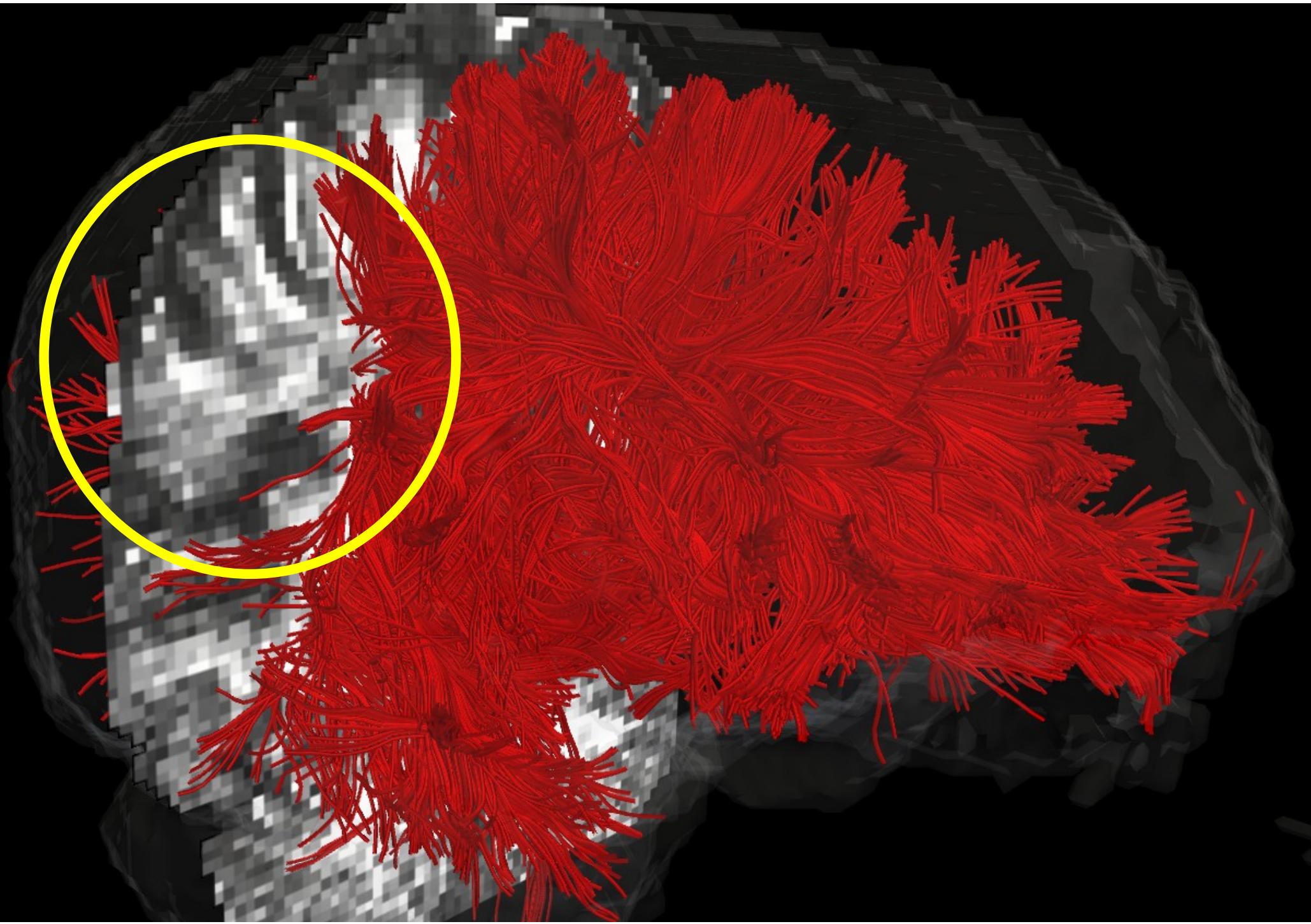
[Jababdi 07]

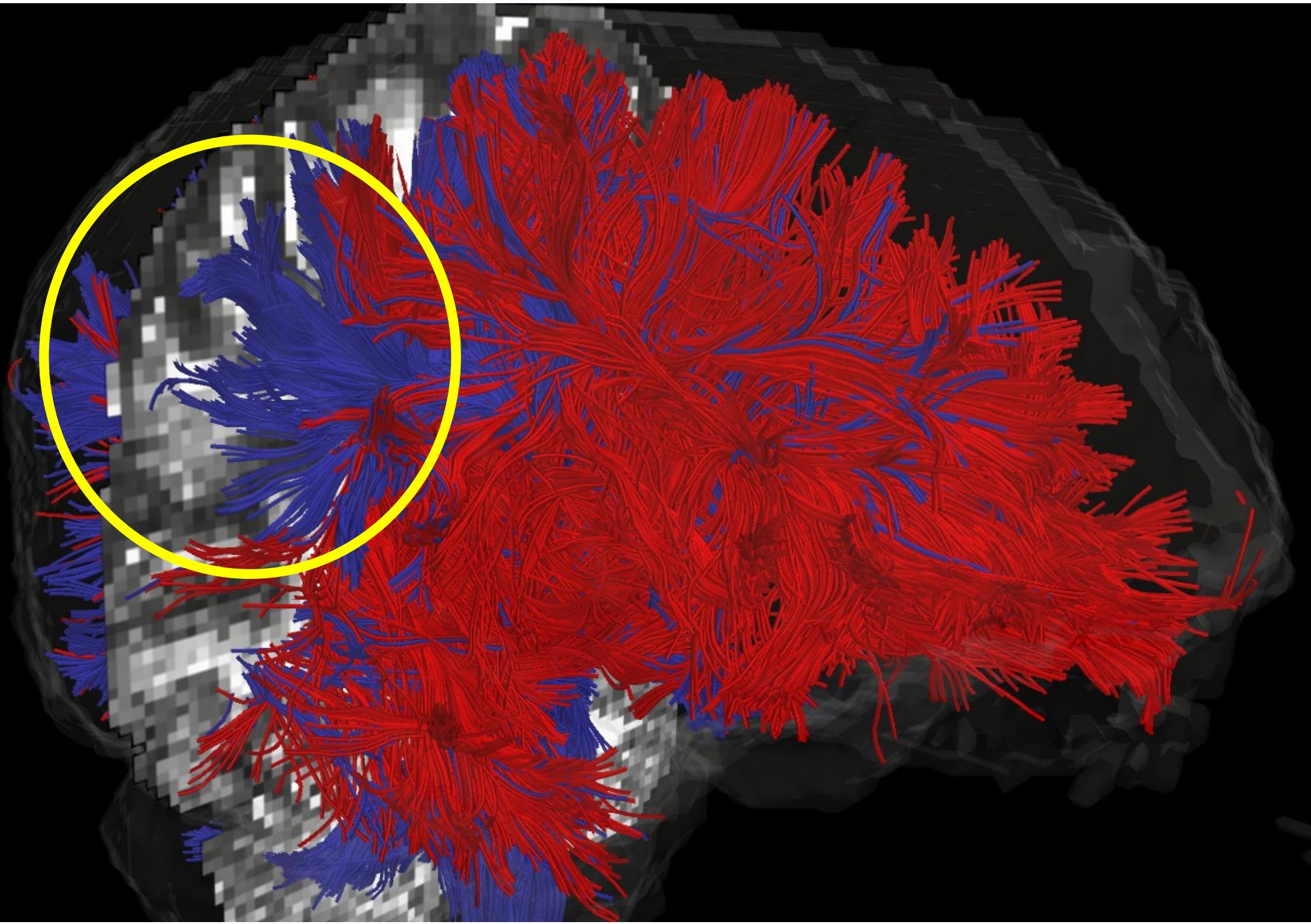
ROI-based  
to constrain  
probabilistic

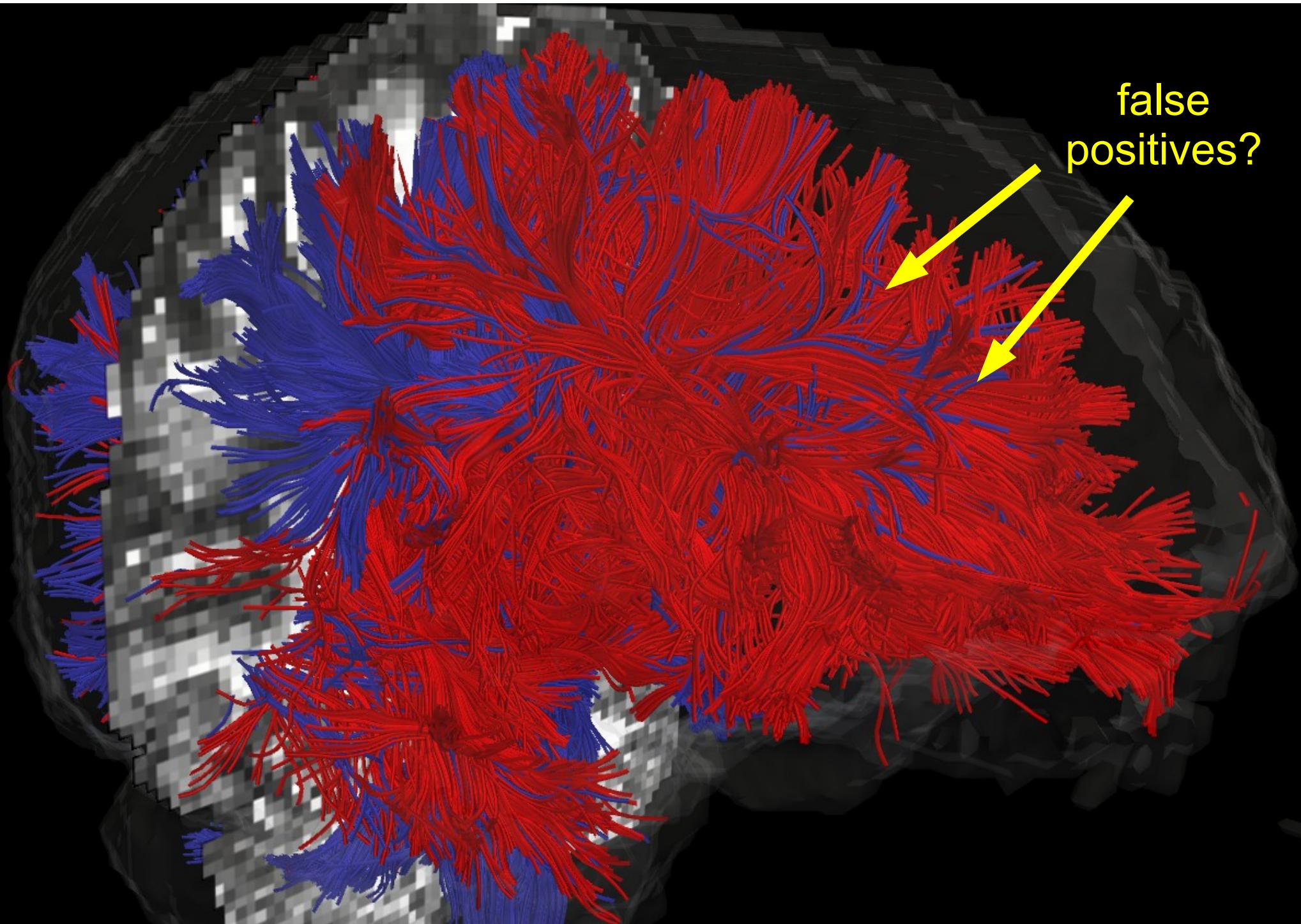


[Poupon 08, Deriche 09]



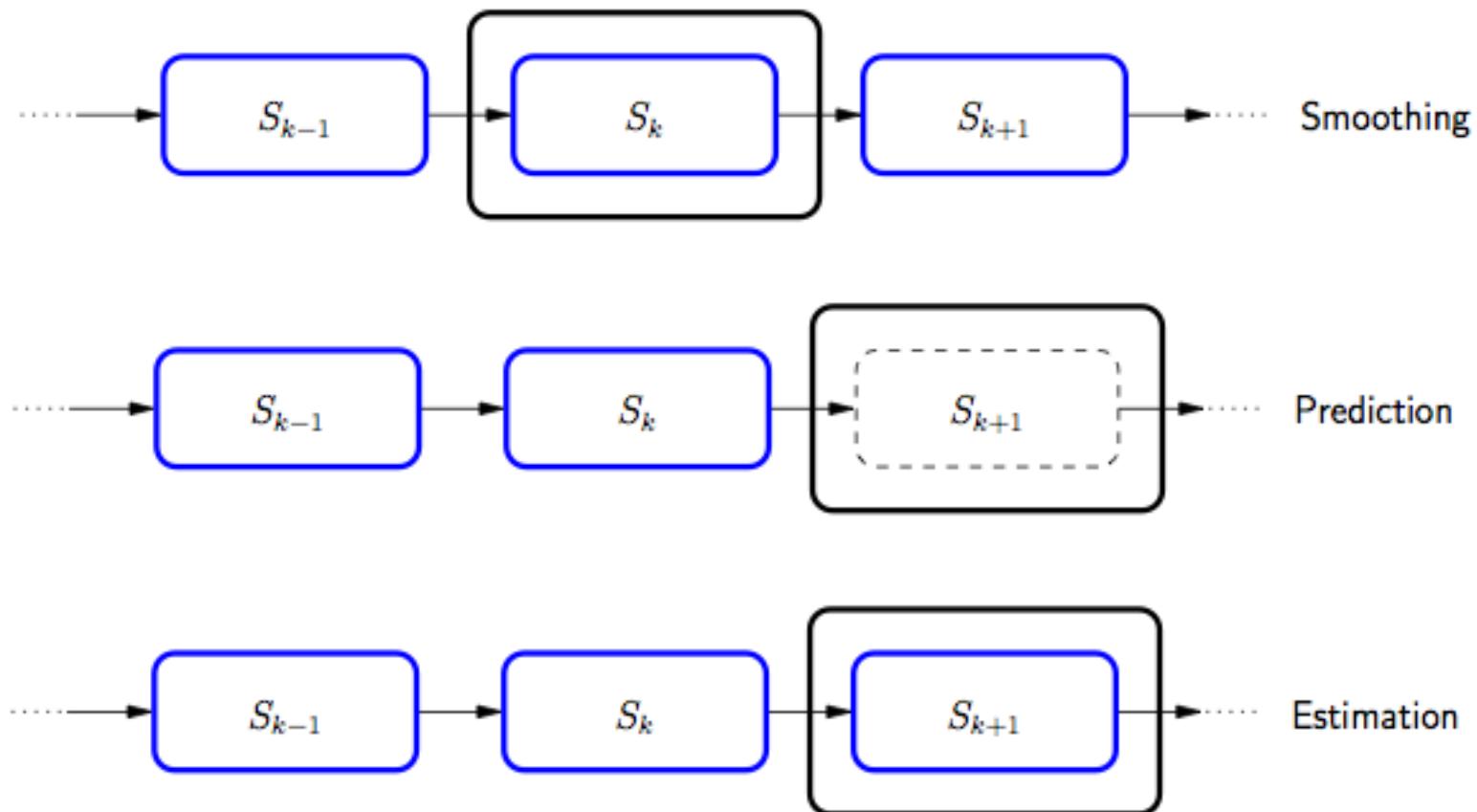






false  
positives?

# filtering



[Niethammer 08]

## brute force optimization

- matching pursuit
- parametric dictionary
- noiseless signal
- discretization, noise

## spherical harmonics

- non-parametric
- order eight (8)
- fiber sharpening for peak detection ( $L=0.006$ )

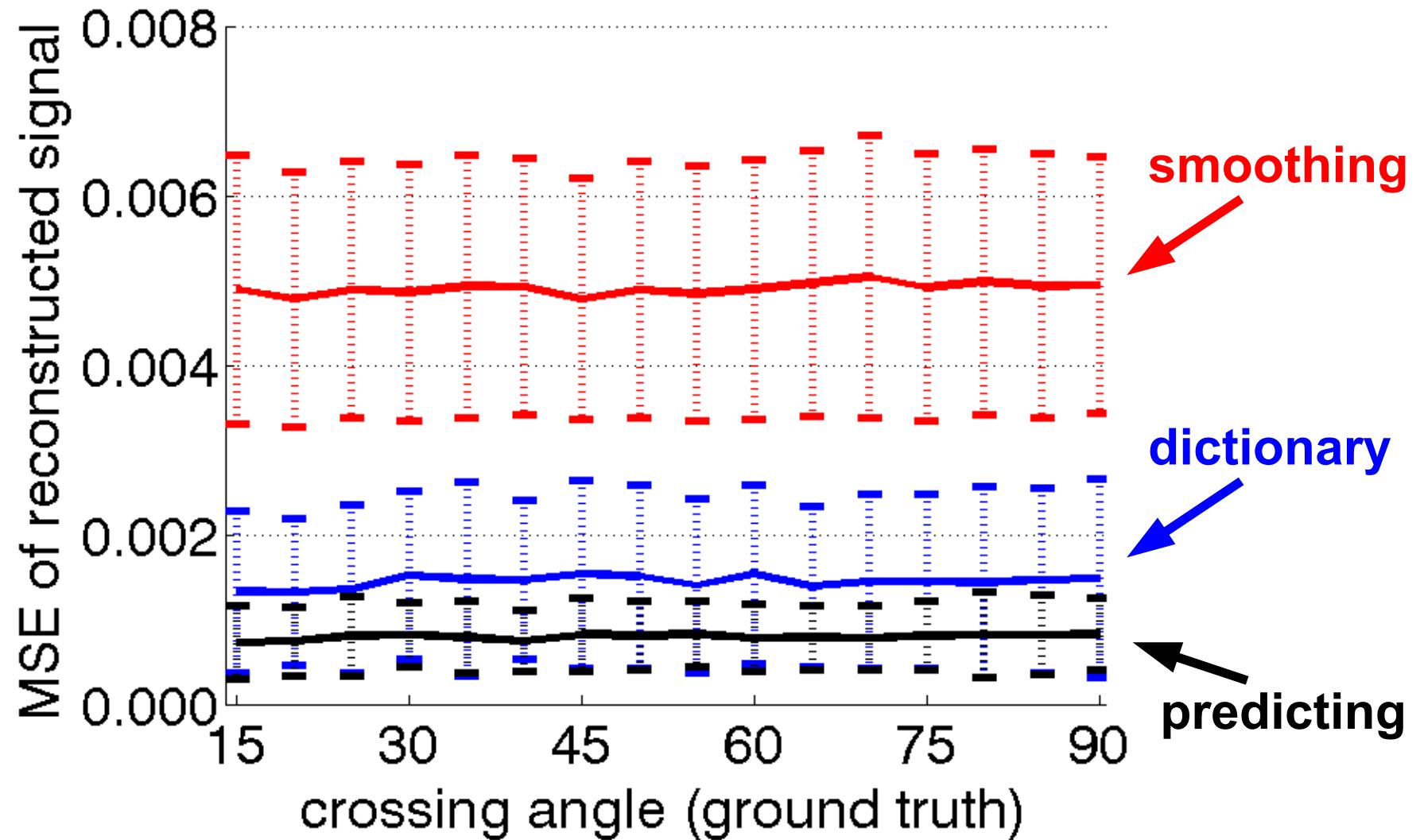
## filtered tractography

- two-fiber model
- unscented Kalman filter

[Descoteaux 07]

J. G. Malcolm, M. E. Shenton, and Y. Rathi. Neural tractography using an unscented Kalman filter. In *Information Processing in Medical Imaging (IPMI)*, pages 126–138, 2009.

# signal reconstruction error



SNR  $\approx 5$ , b = 1000