



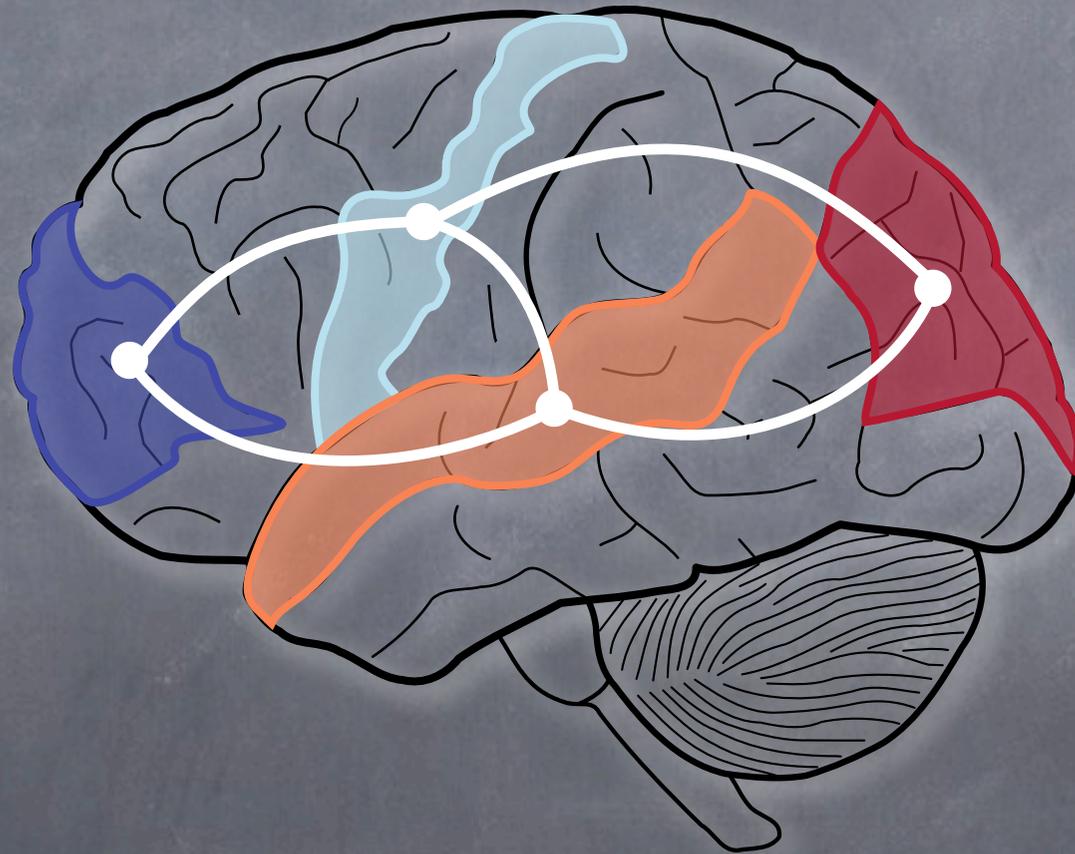
Multi-Fascicle Models for Population Studies of the Brain Microstructure

Soutenance de thèse
Louvain-la-Neuve, October 2013

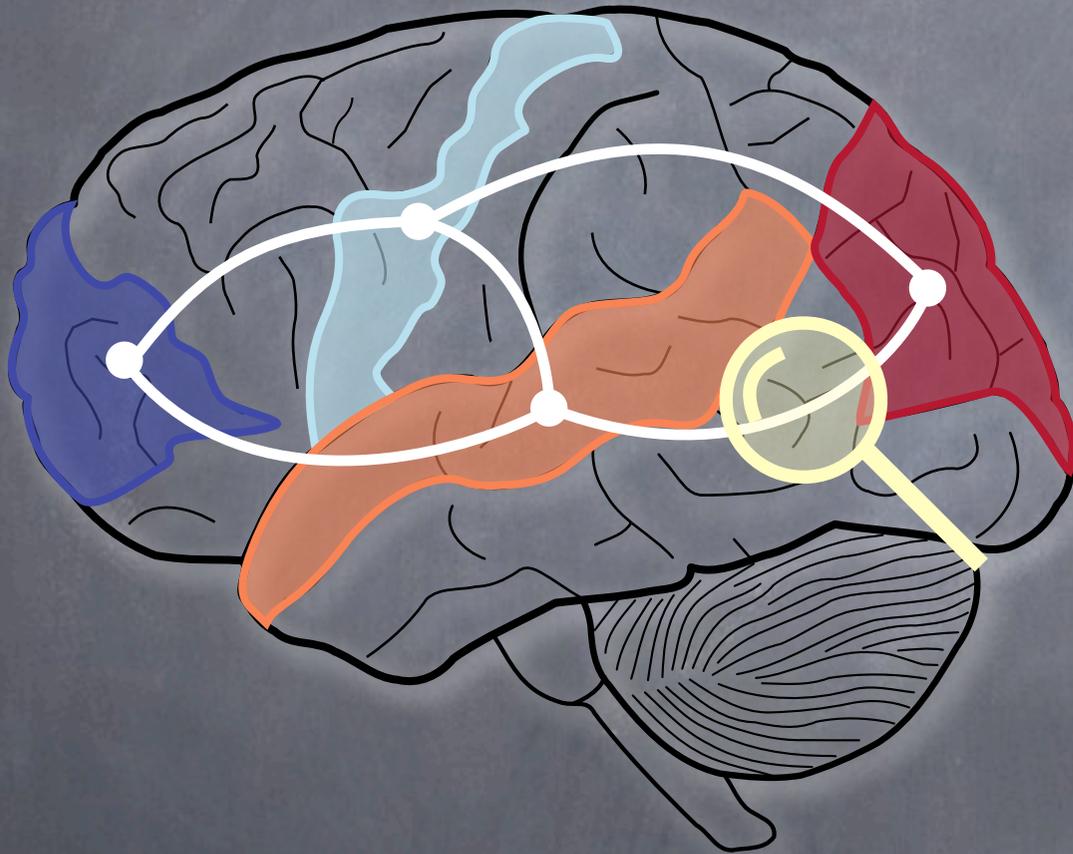
Maxime Taquet



Disconnection syndromes and other disorders are thought to affect the connections in the brain



The microstructure underpins connectivity



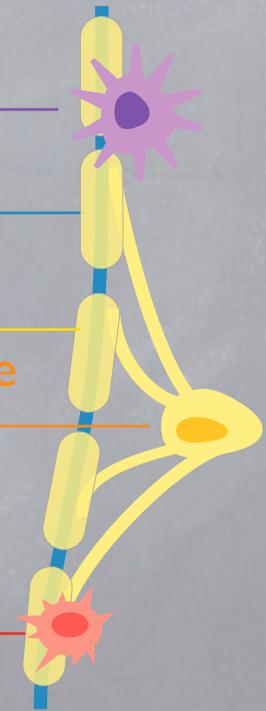
Astrocyte

Axon

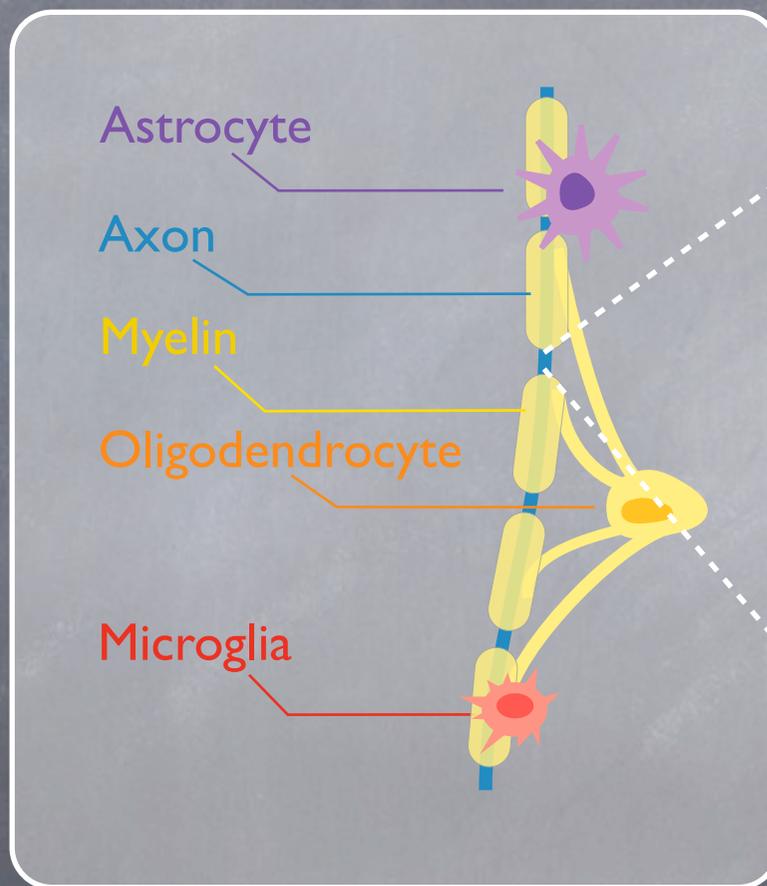
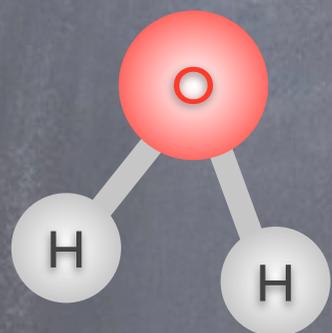
Myelin

Oligodendrocyte

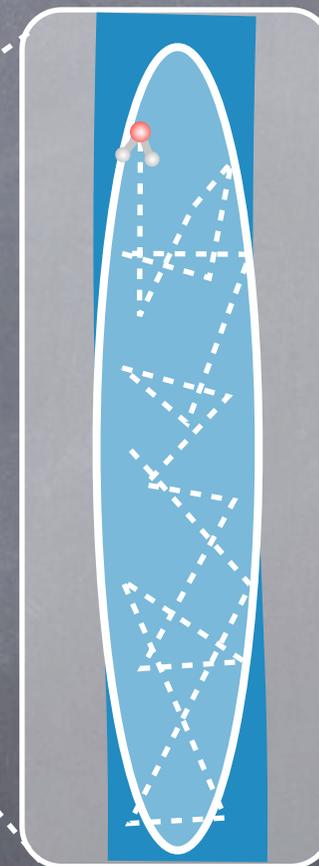
Microglia



Diffusion imaging can probe the microstructure

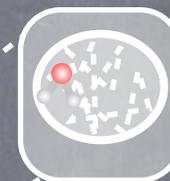
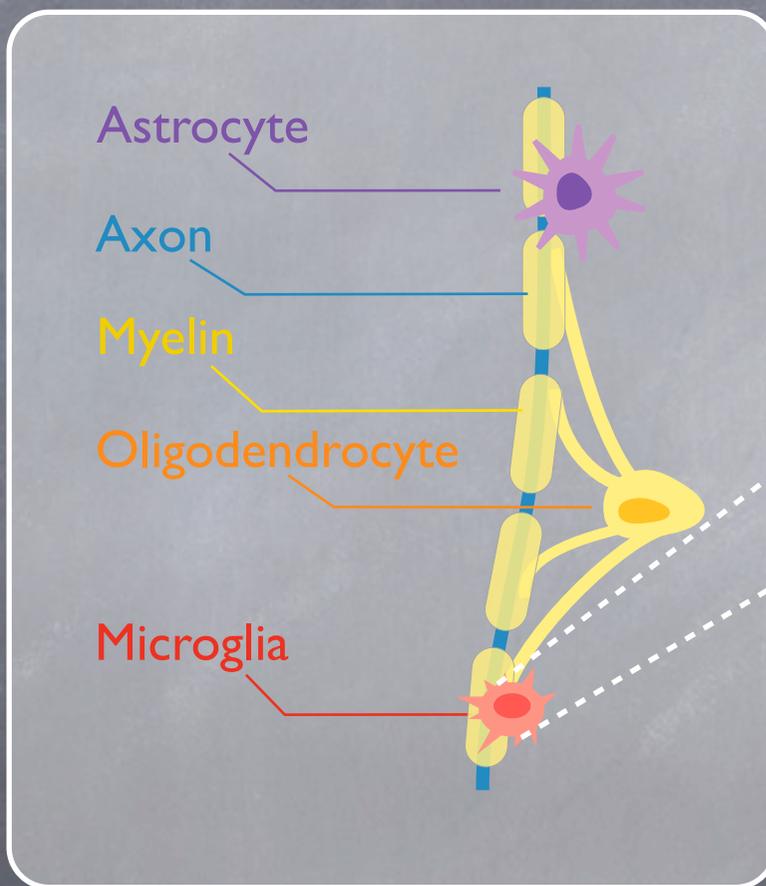
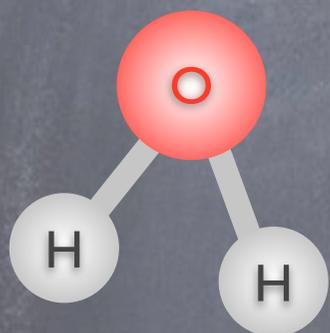


Anisotropic
restricted diffusion

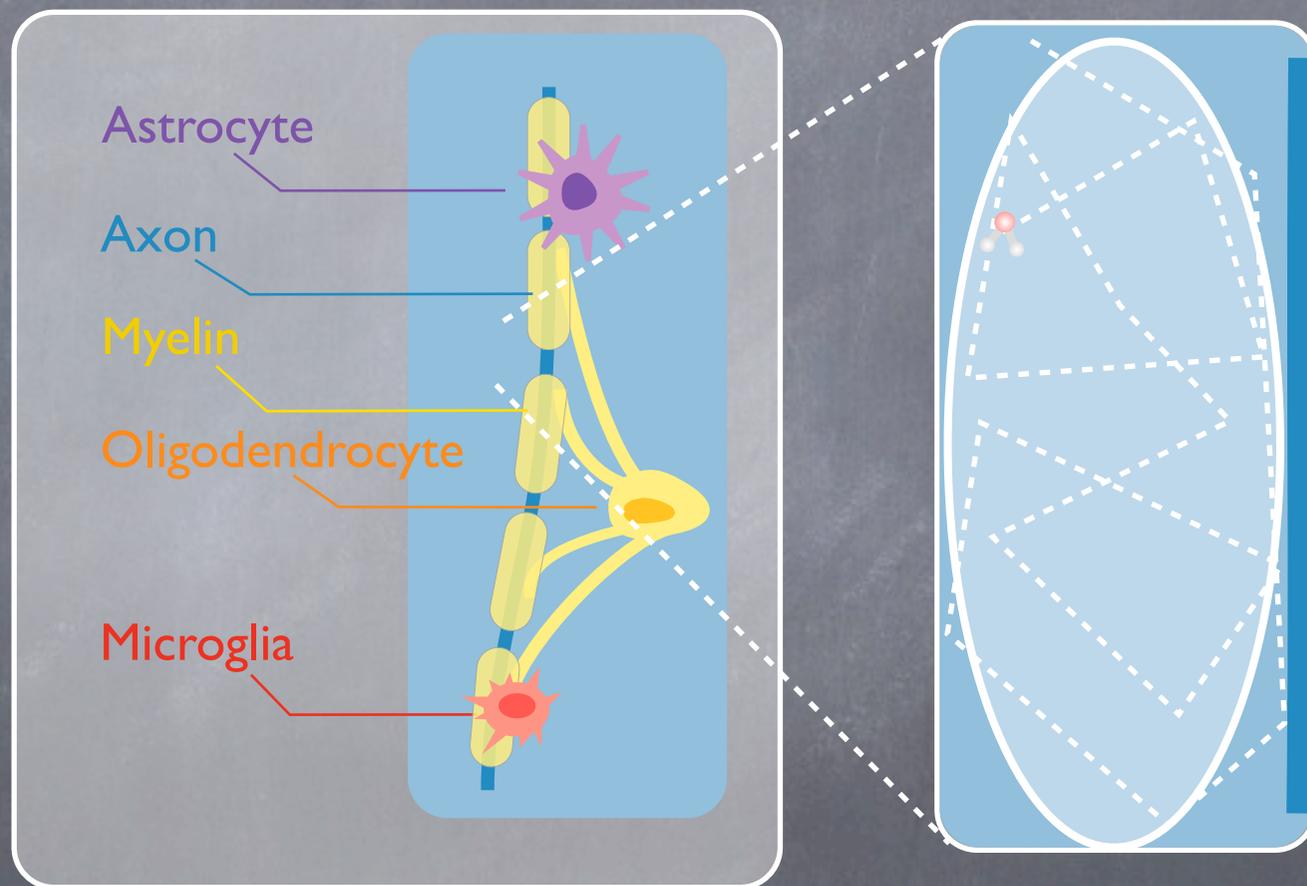
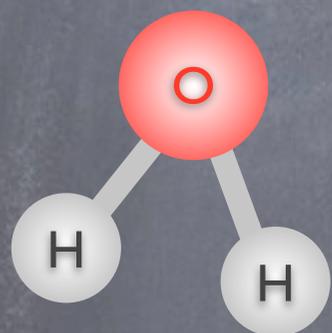


Diffusion imaging can probe the microstructure

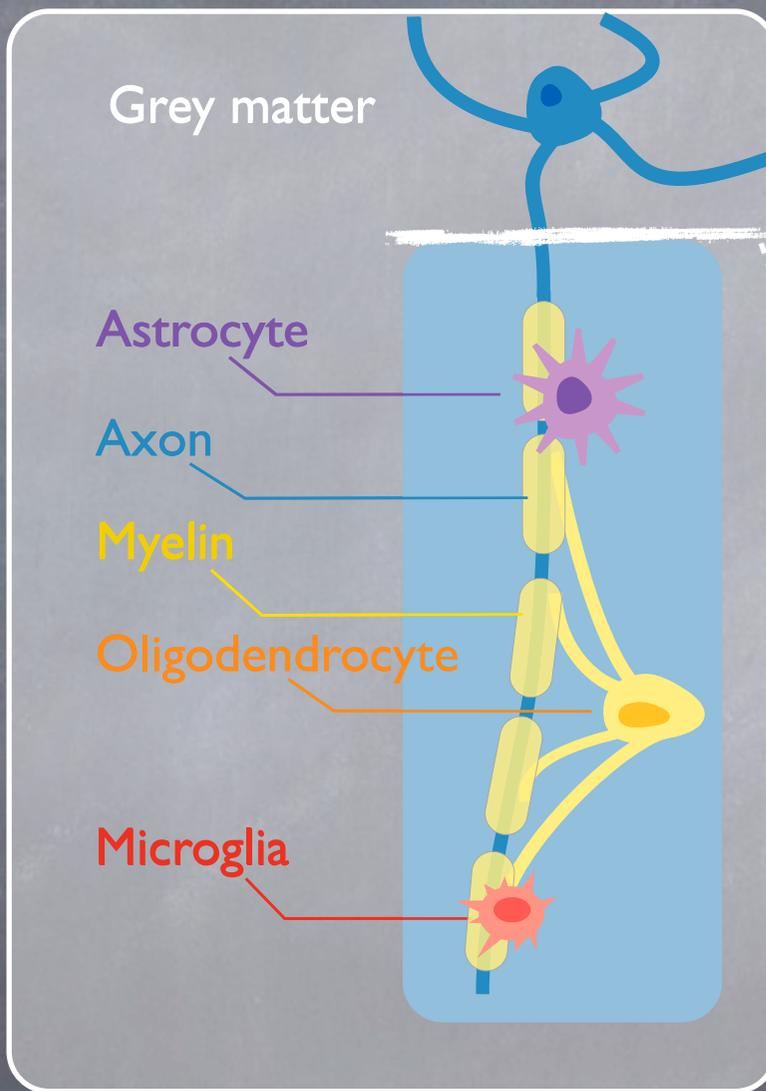
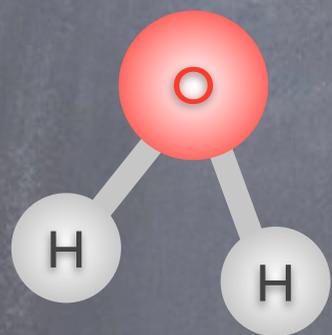
Isotropic
restricted diffusion



Diffusion imaging can probe the microstructure



Diffusion imaging can probe the microstructure

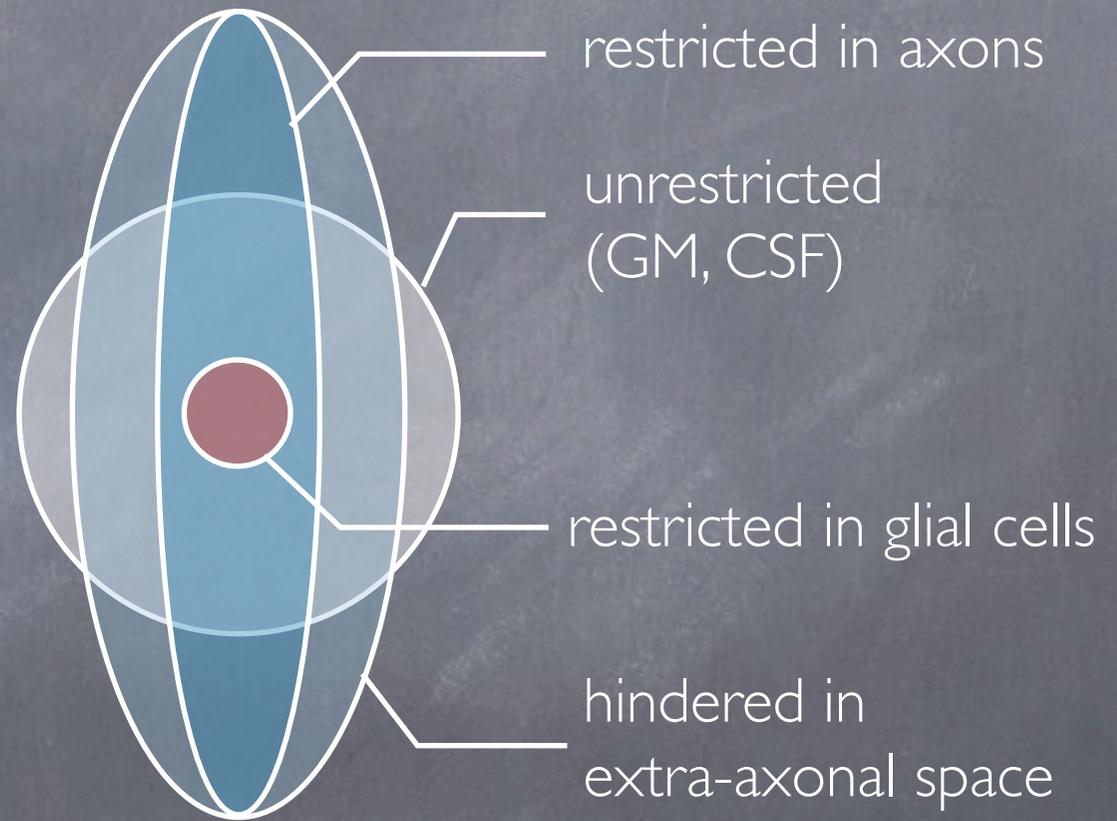
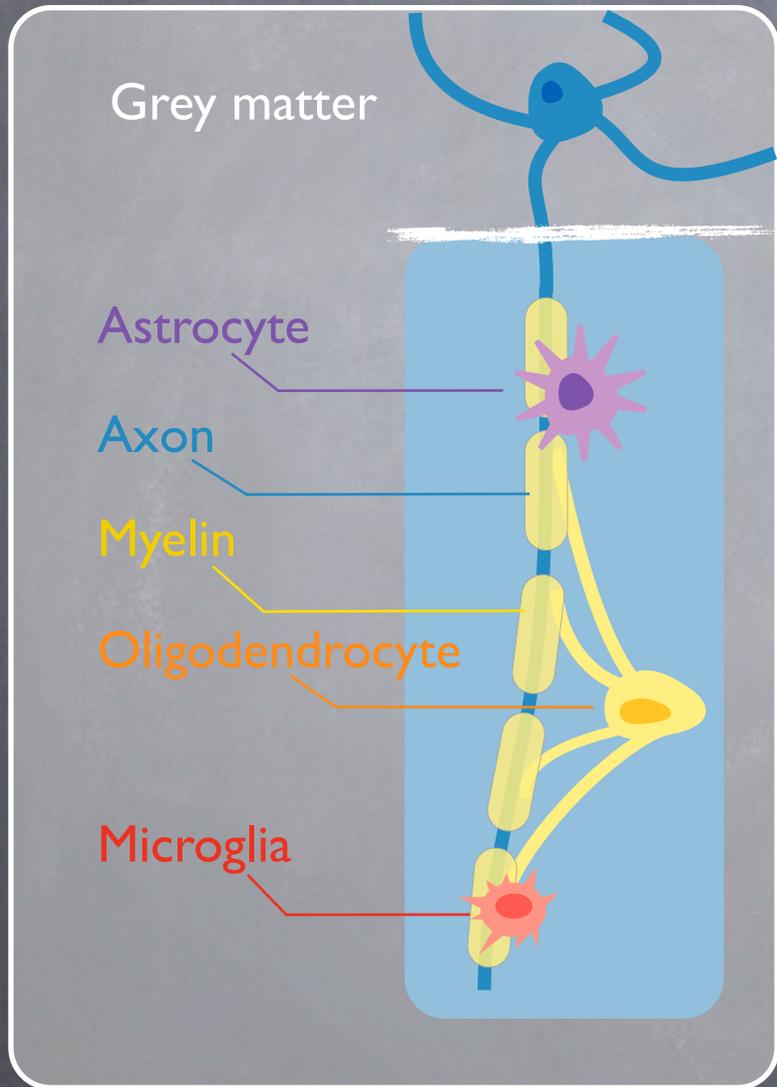


Isotropic
unrestricted diffusion

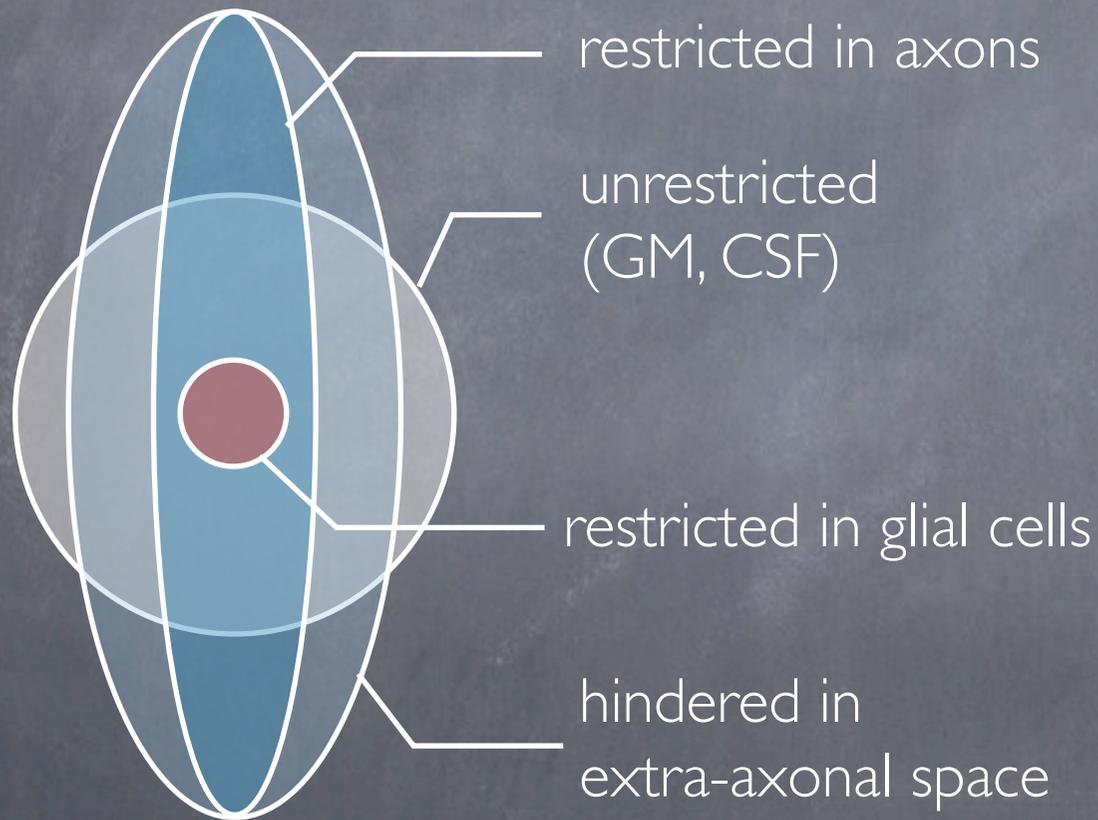


Diffusion imaging can probe the microstructure

If there is no exchange of water molecules between the different compartments



Diffusion imaging can probe the microstructure

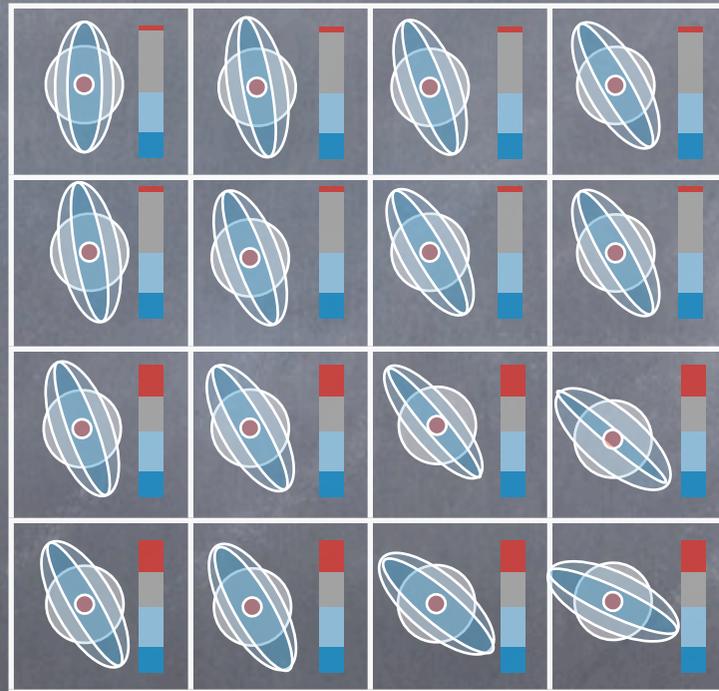


Fraction of water molecules in each compartment

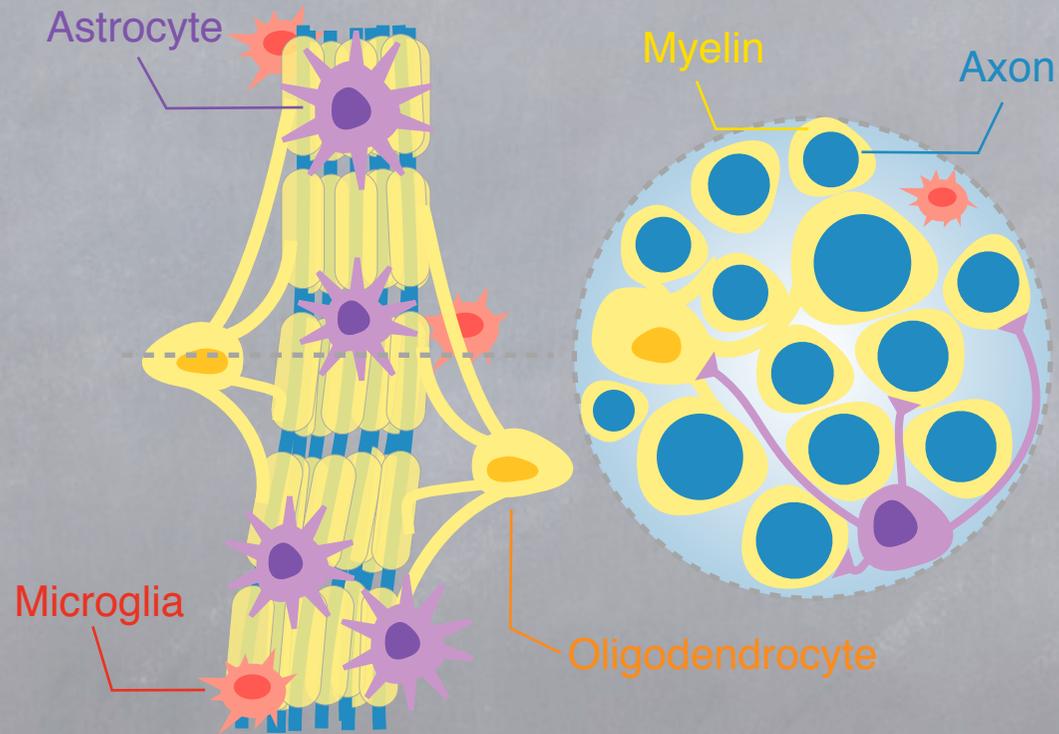


Images of the brain microstructure contain a model in each voxel

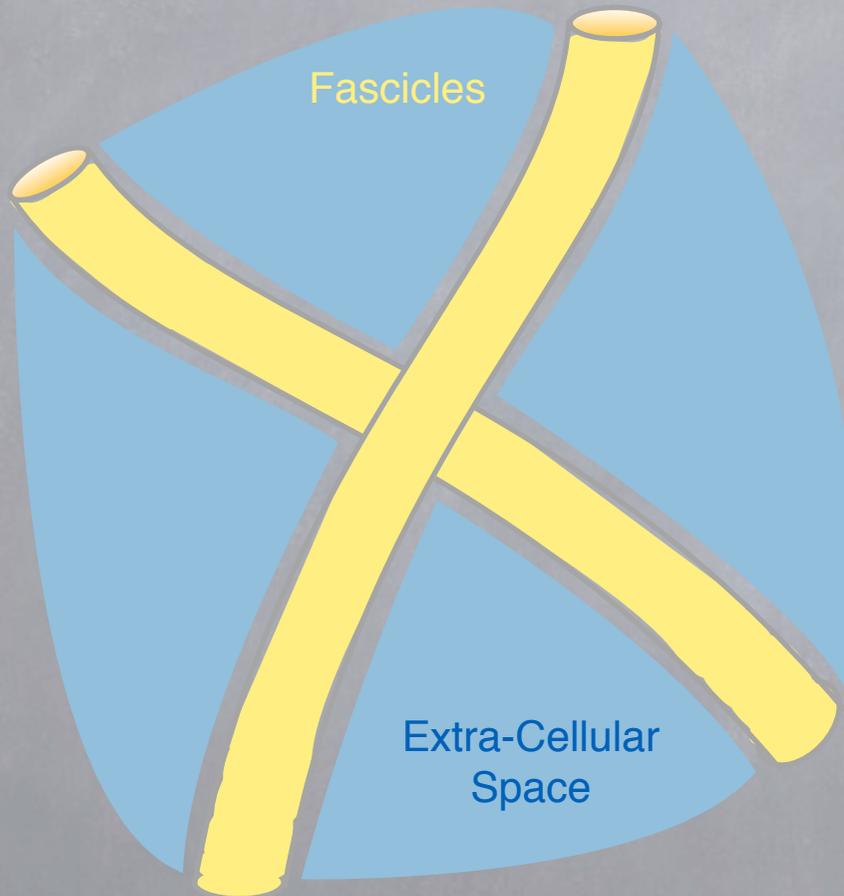
The models in different voxels have different parameters



Axons travel in fascicles



In most voxels, several fascicles cross each other

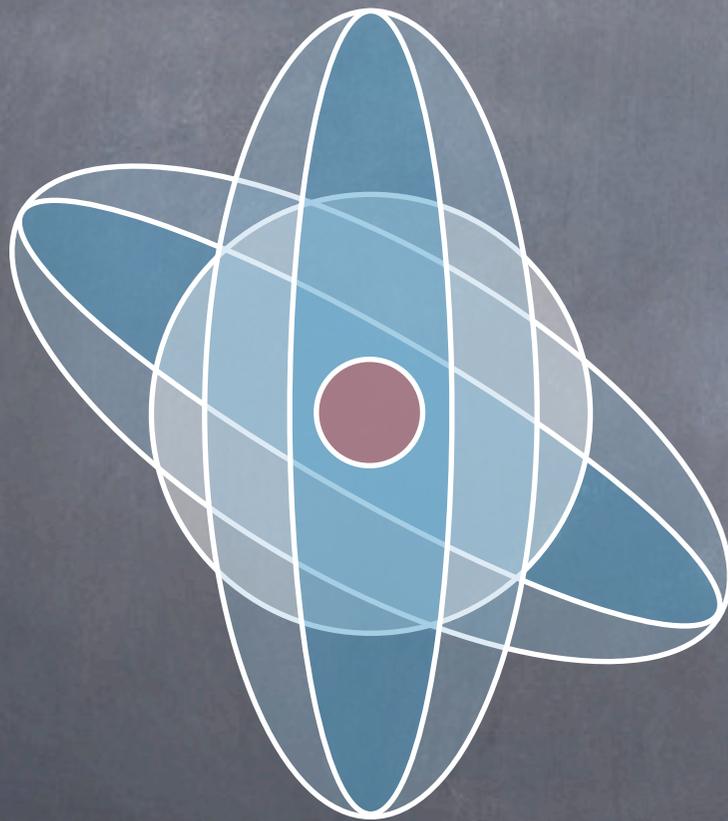


At typical resolution,
60-90% of the brain voxels
contain crossing fascicles

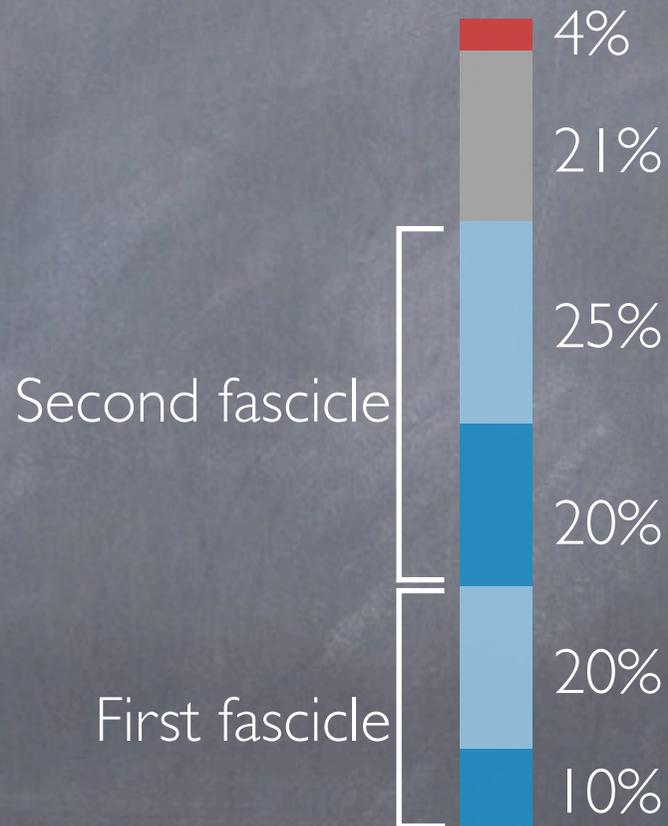
Jeurissen et al., 2012

In most voxels, several fascicles cross each other

Multi-Fascicle models of the brain microstructure

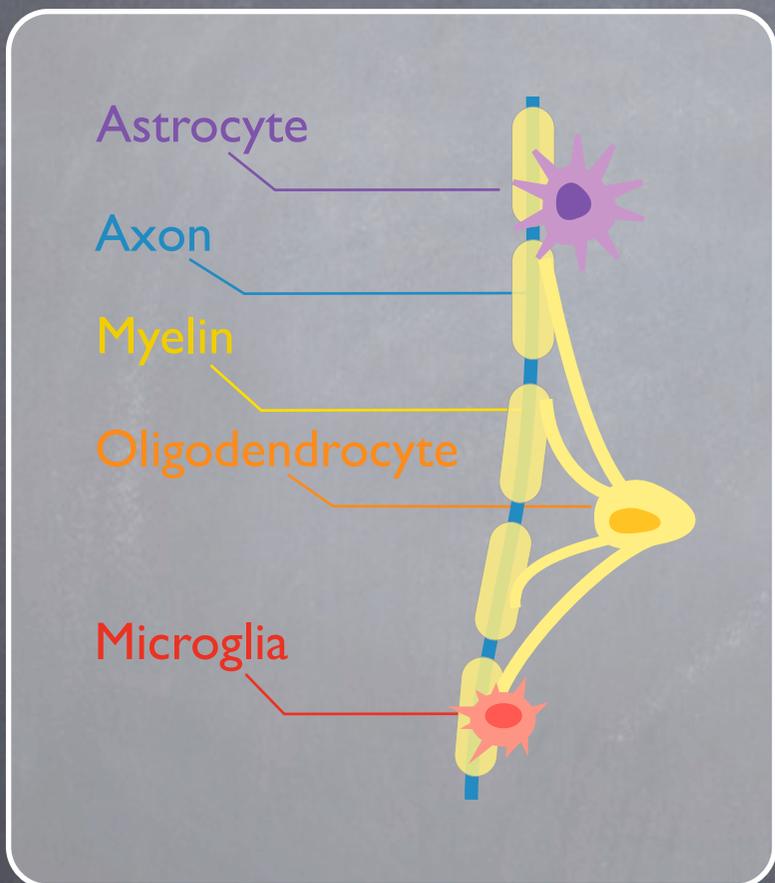


Fraction of water molecules in each compartment

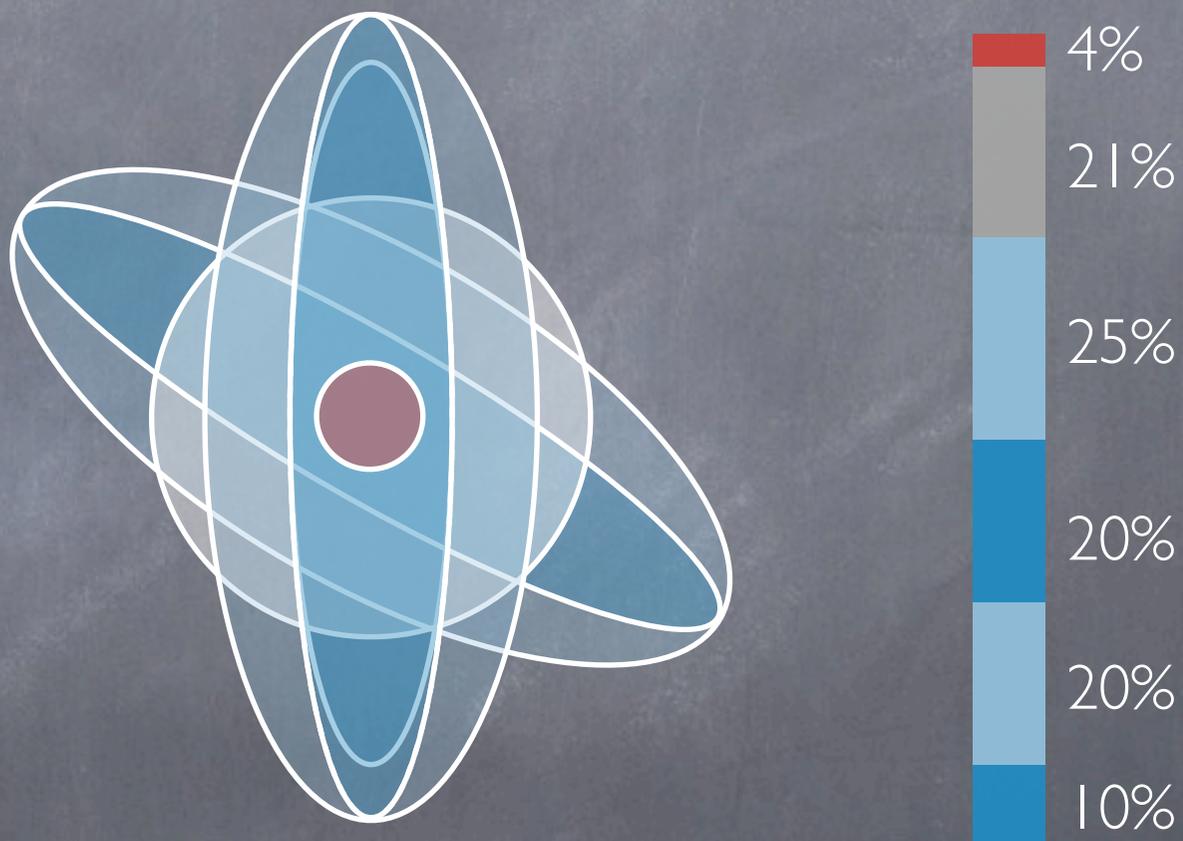


Myelin injury is translated in a decreased axial diffusivity

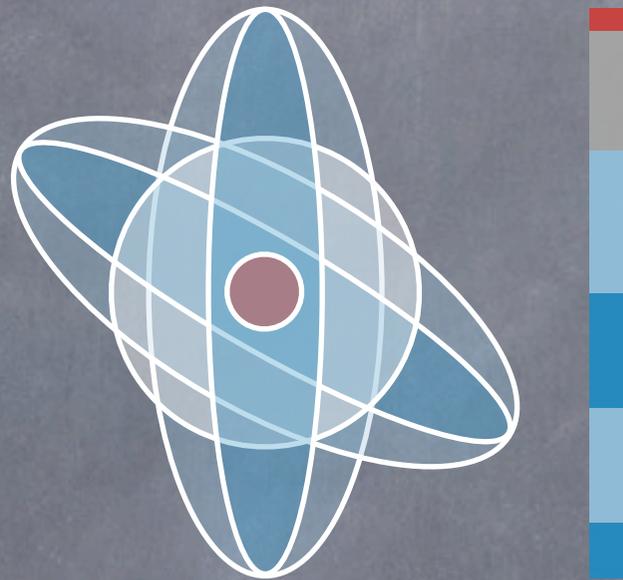
Axon injury



Multi-Fascicle models of the brain microstructure



Population studies of the brain microstructure from diffusion imaging



Acquire
Diffusion-Weighted
MRI

Estimate
Microstructural
Models

Register
Microstructural
Models

Perform
Statistical Analysis

New assets and capabilities to conduct population studies of the brain microstructure

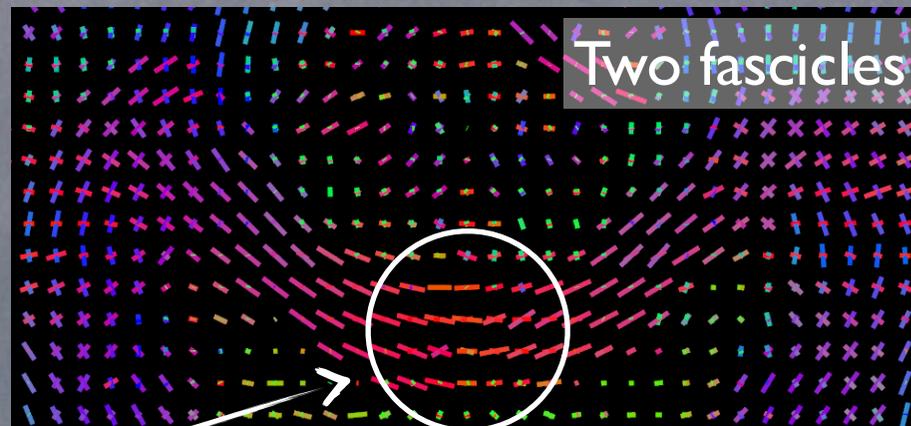
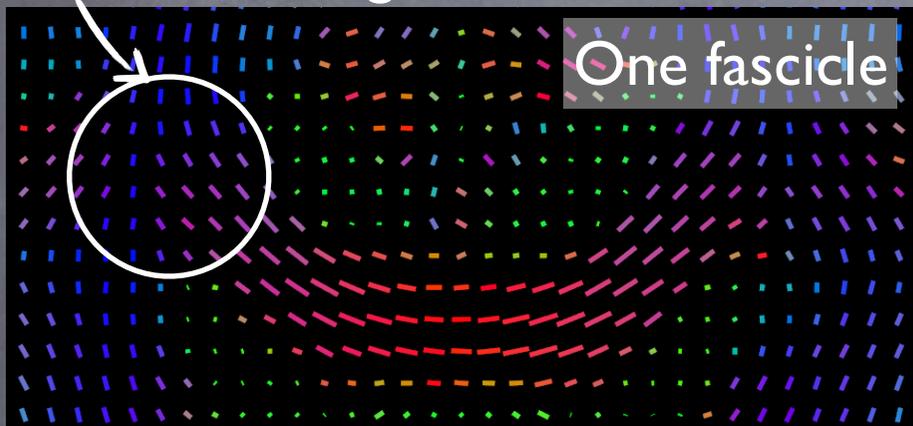
1. Selection of the appropriate model
2. Registration and atlas construction
3. Statistical analysis of microstructure
4. Estimation from single b-value data

New assets and capabilities to conduct population studies of the brain microstructure

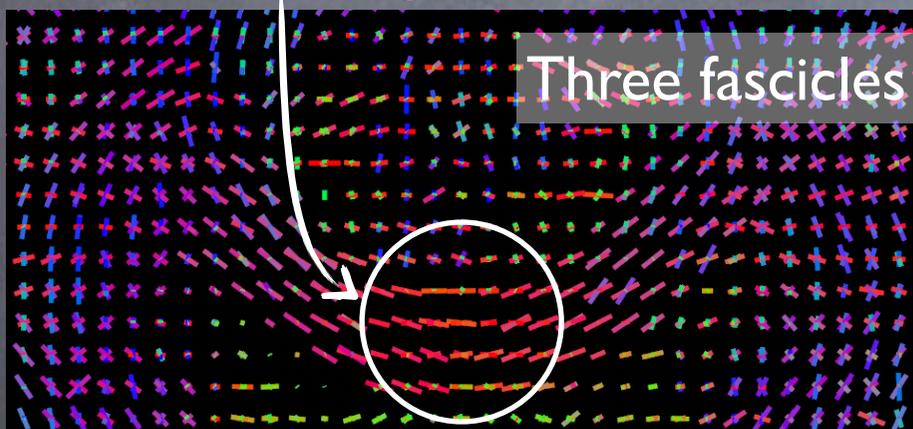
1. Selection of the appropriate model
2. Registration and atlas construction
3. Statistical analysis of microstructure
4. Estimation from single b-value data

Given a set of data, we ought to know what model can be reliably estimated

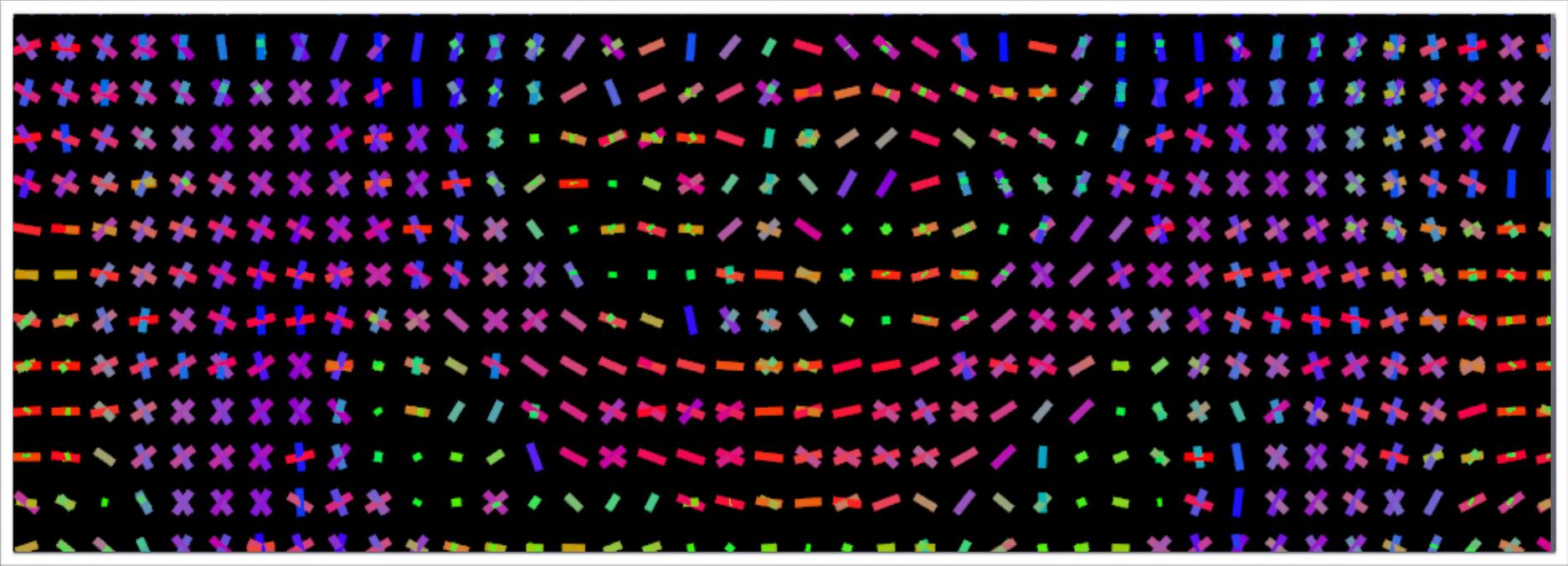
Misses crossing fascicles



Overfitting



Given a set of data, we ought to know
what model can be reliably estimated

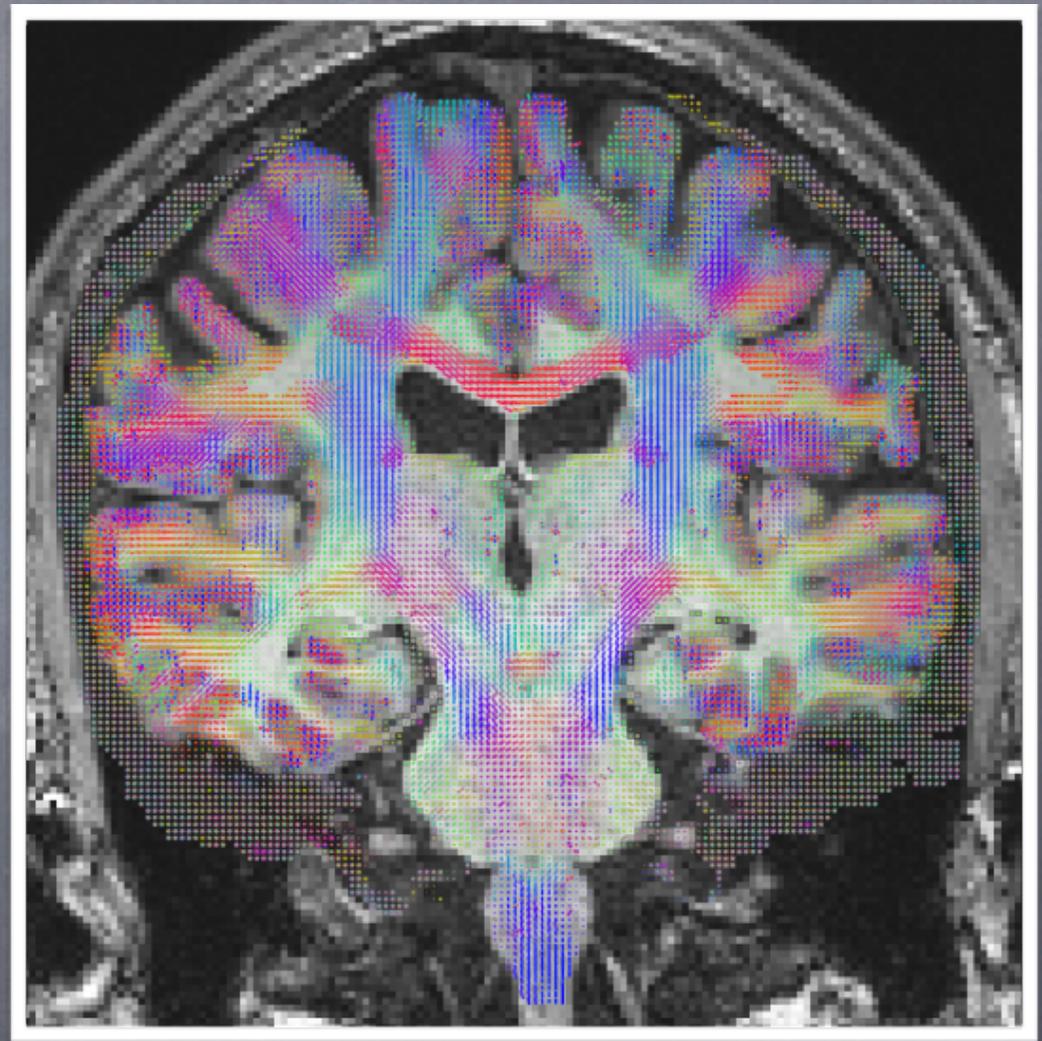
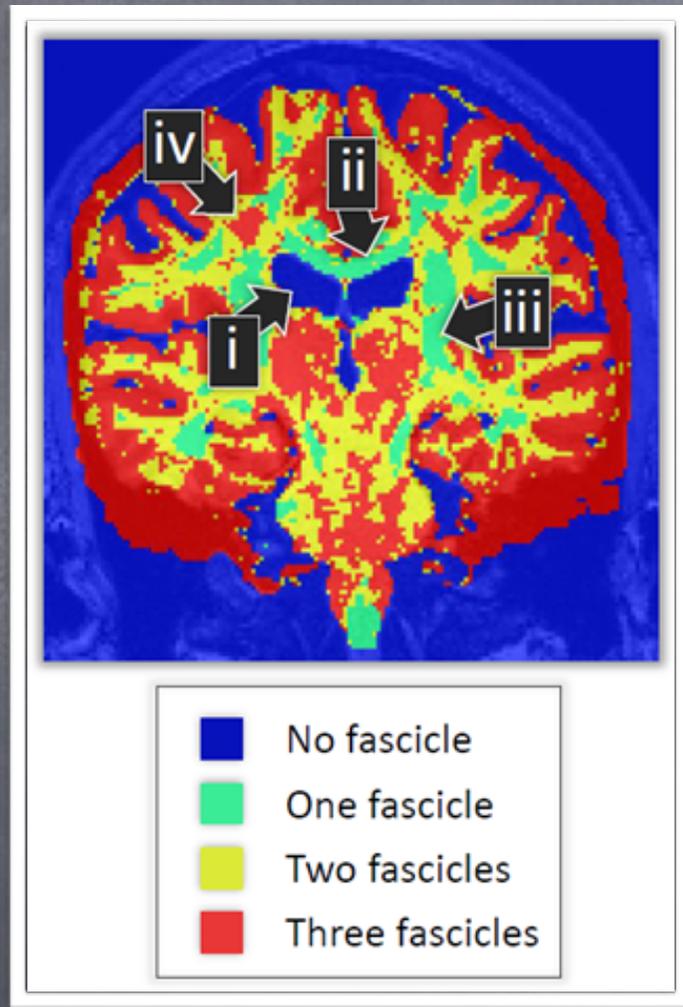


We proposed that the appropriate model best generalizes to new data not included in the estimation

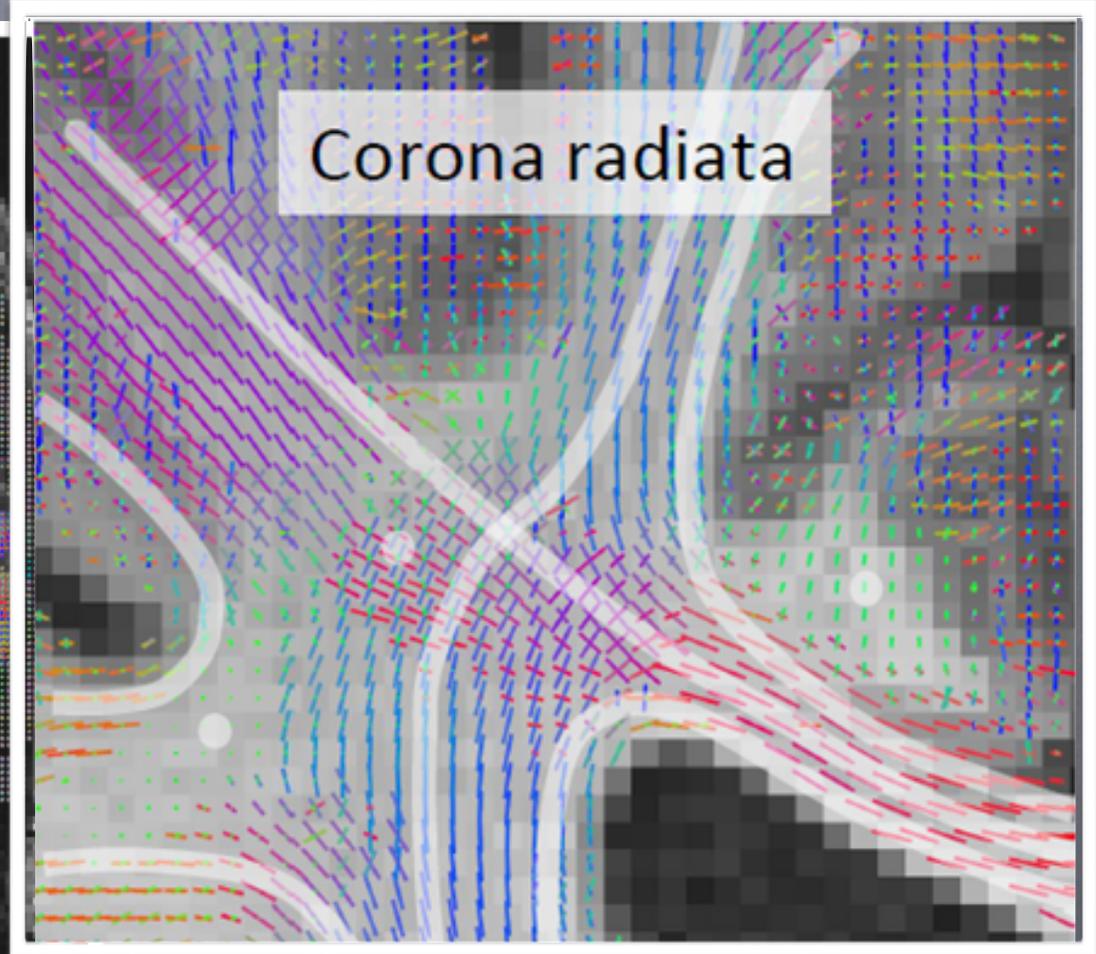
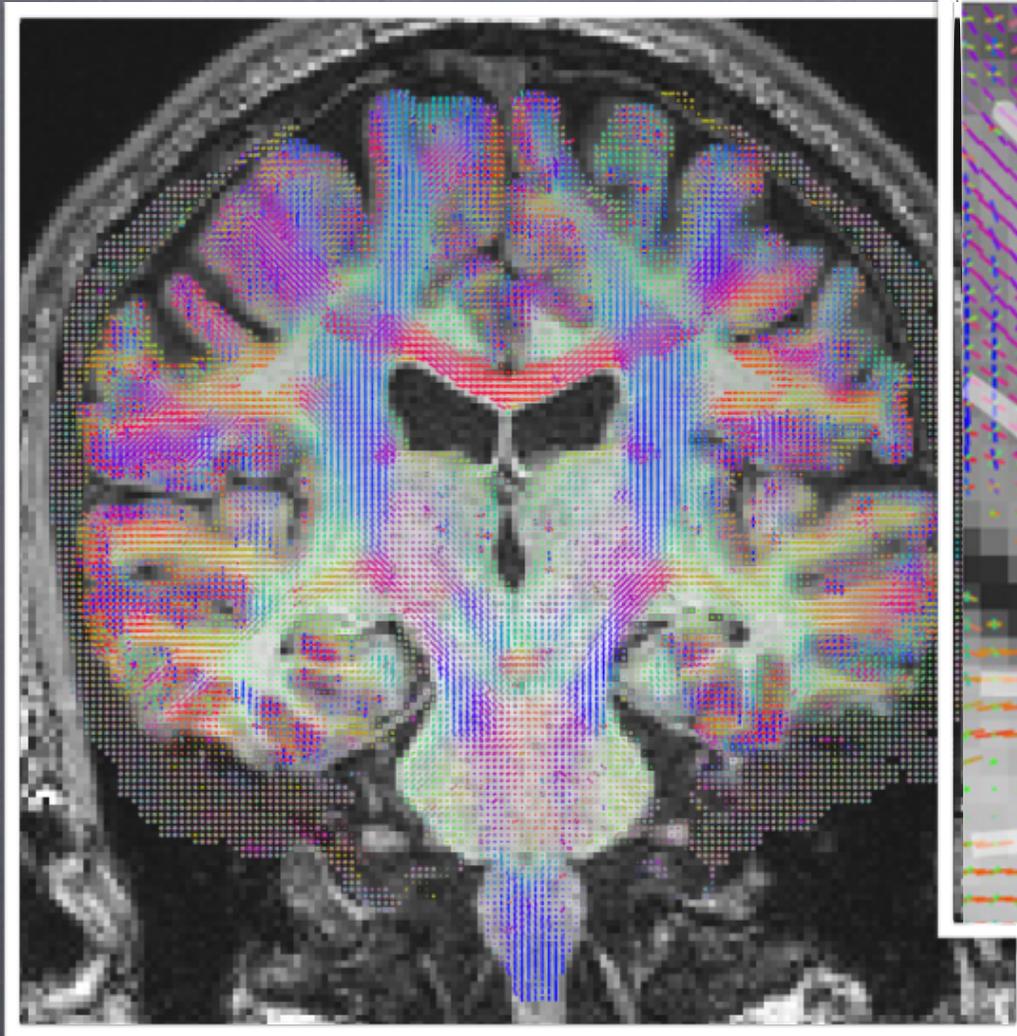


→ Compute the generalization error
632 Bootstrap estimator

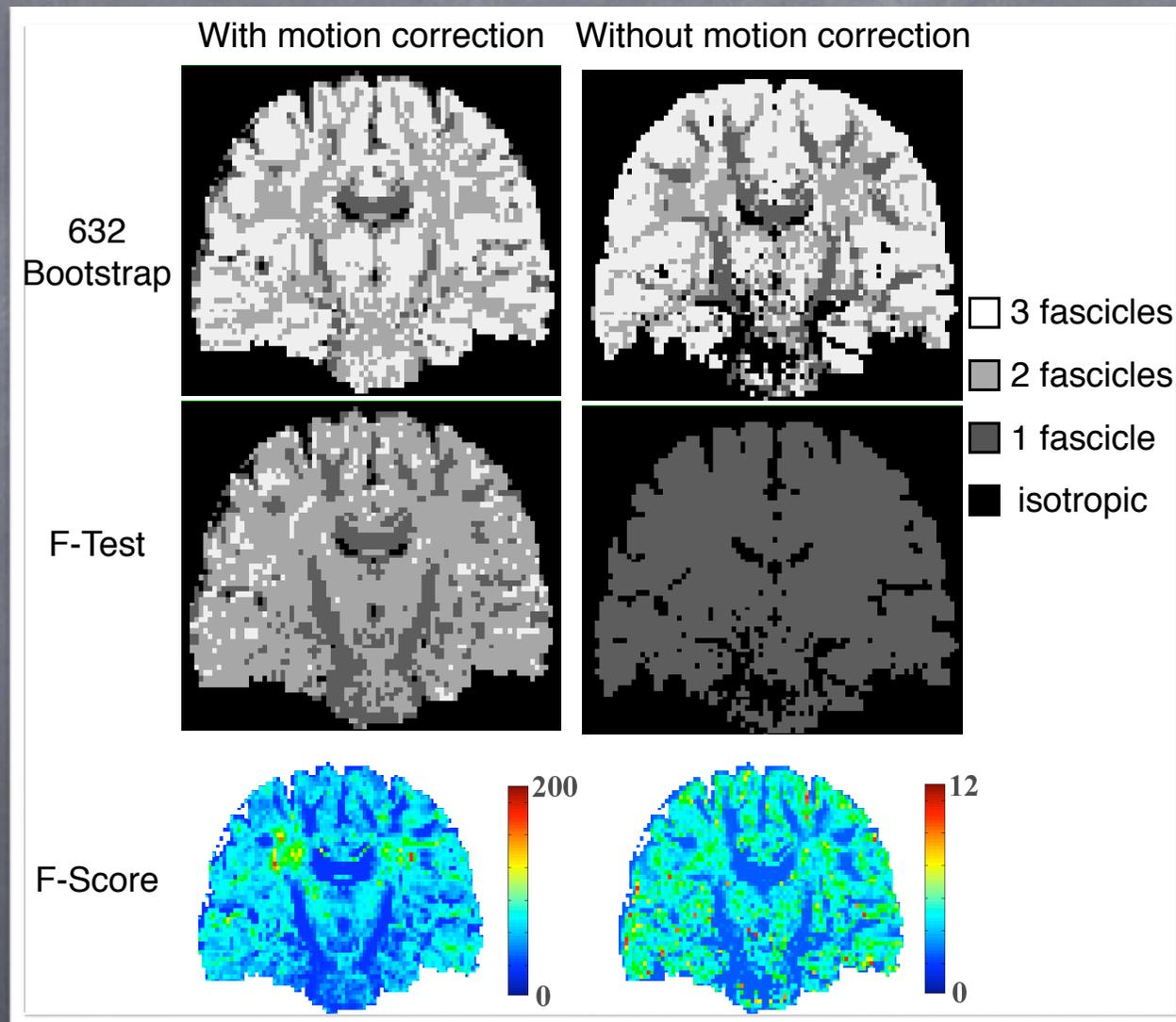
As a result, we obtain a reliable and robust selection of the microstructure model



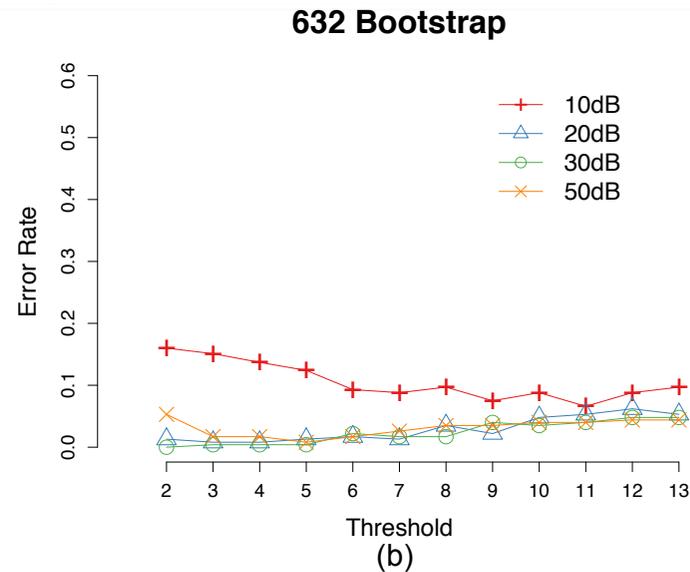
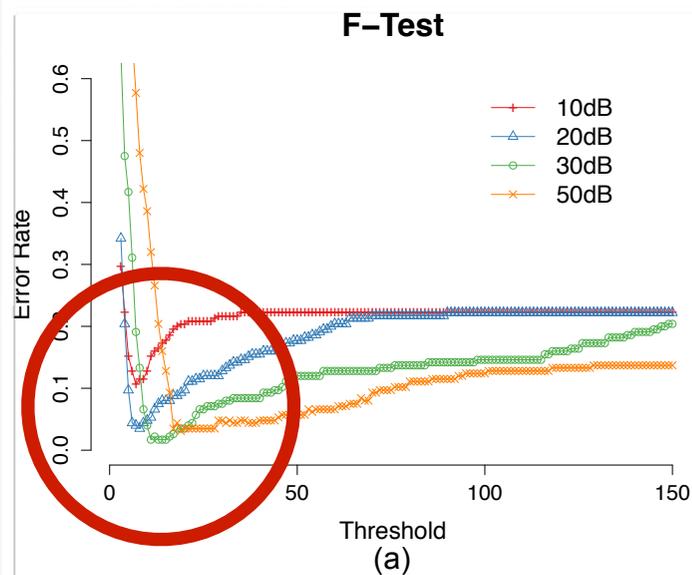
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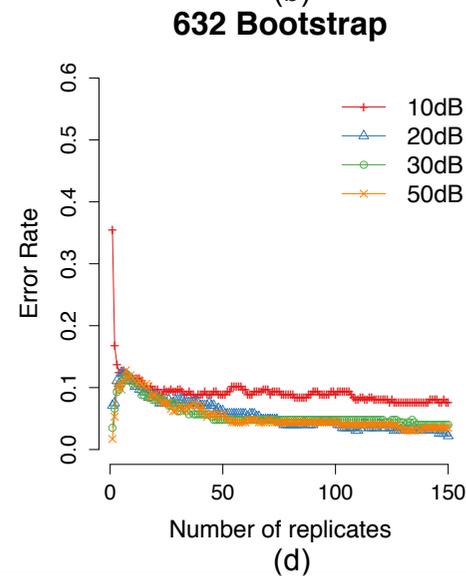


As a result, we obtain a reliable and robust selection of the microstructure model



		10dB	20dB	30dB	50dB
overall minimum	F-Test	24 (10%)	8 (3%)	4 (2%)	7 (3%)
	632 Bootstrap	15 (7%)	2 (0.9%)	0 (0%)	2 (0.9%)
single threshold	F-Test	44 (19%)	20 (9%)	6 (3%)	8 (4%)
	632 Bootstrap	20 (9%)	3 (1%)	4 (2%)	6 (3%)

(c)



New assets and capabilities to conduct population studies of the brain microstructure

1. Selection of the appropriate model
2. Registration and atlas construction
3. Statistical analysis of microstructure
4. Estimation from single b-value data

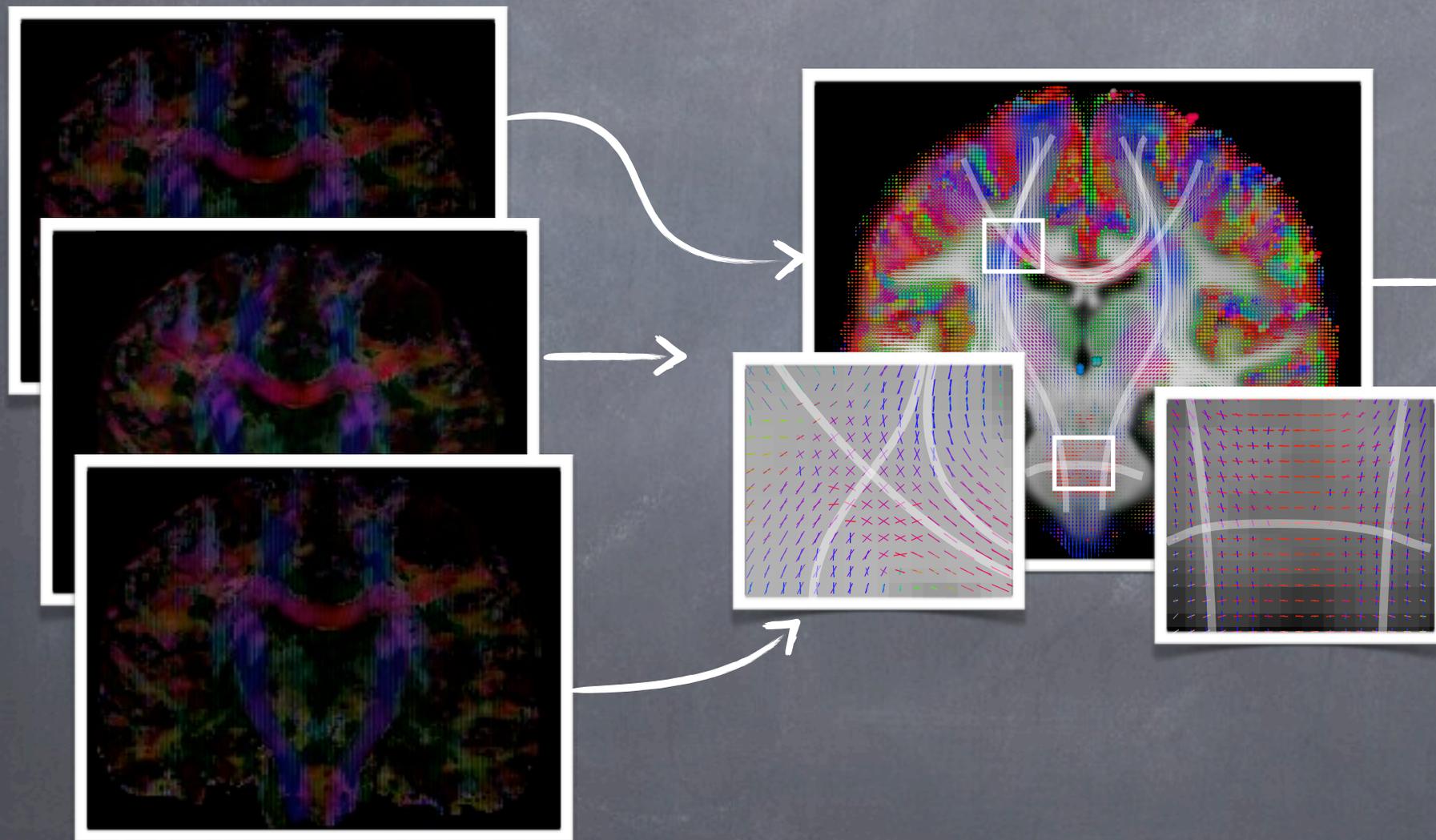
Registration and atlas construction are at the heart of population studies

Population

Registration

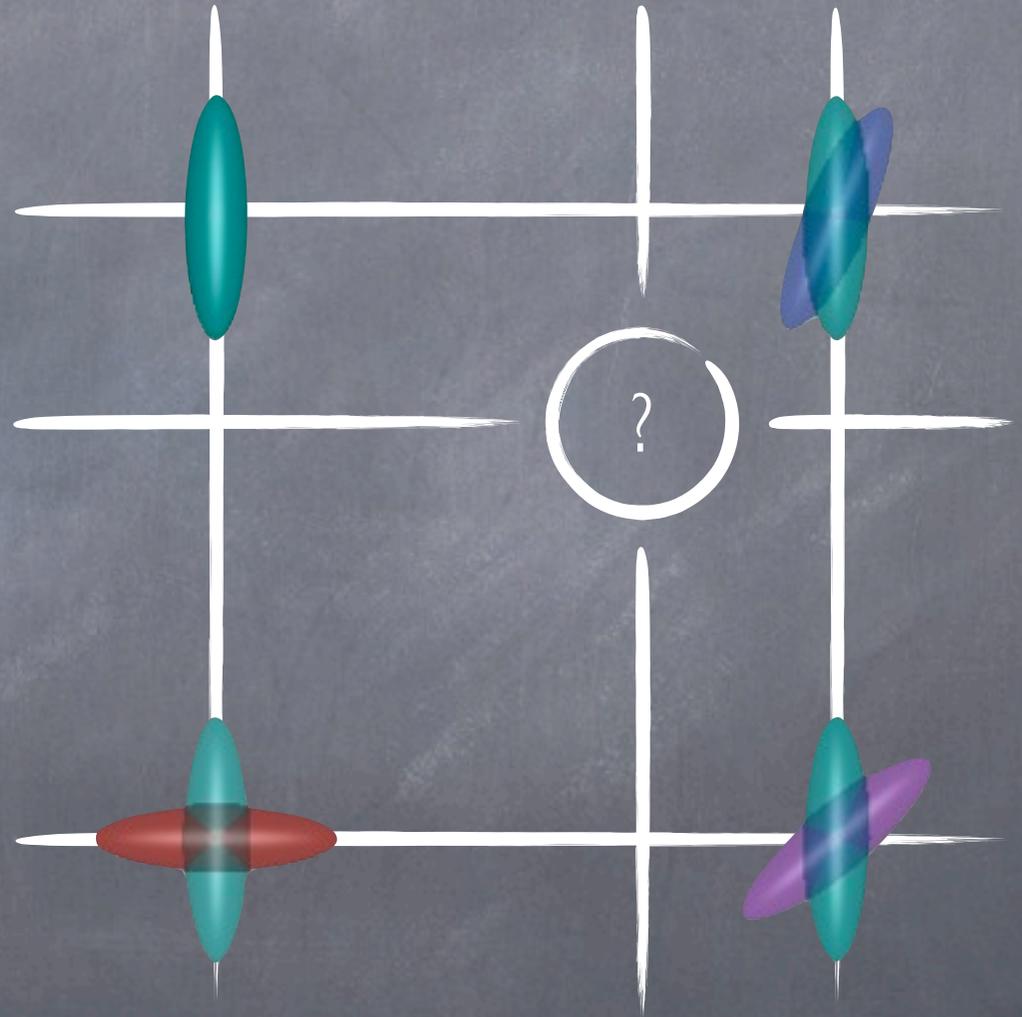
Atlas

Statistical
Analysis



Registration and atlasing require specific operations

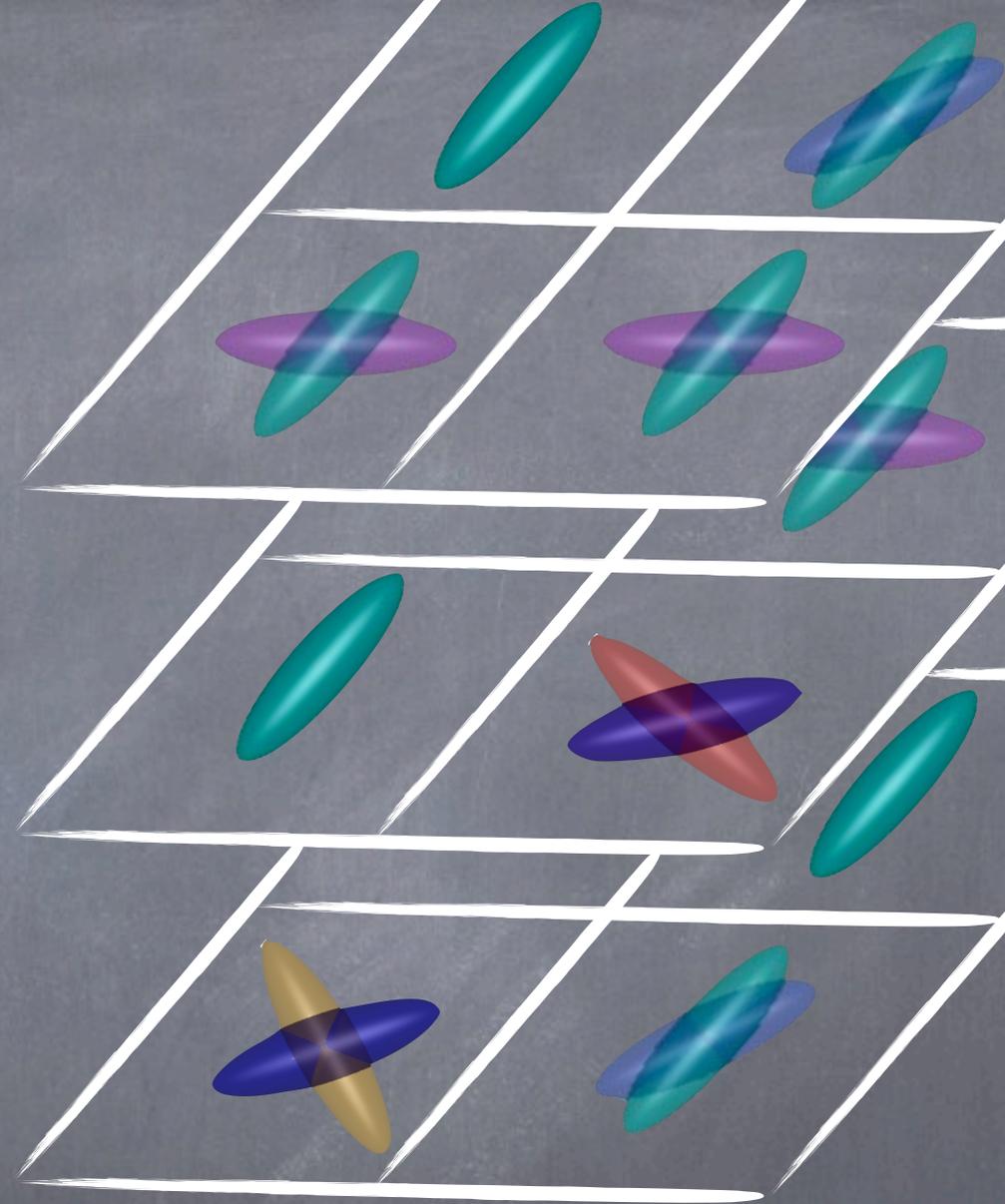
Interpolation



Registration and atlasing require specific operations

Interpolation

Averaging



Registration and atlasing require specific operations

Interpolation
Averaging

$1/3$

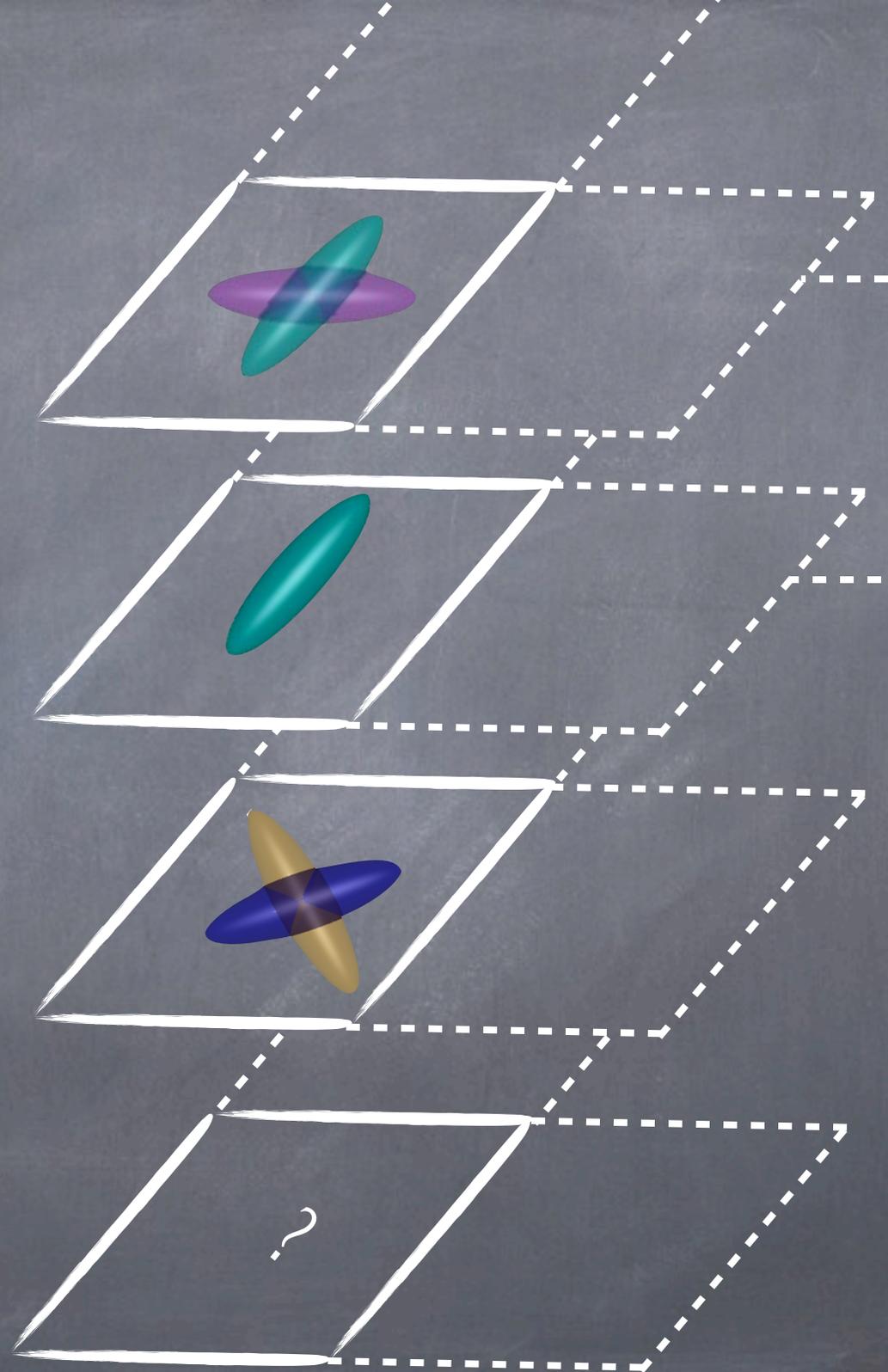
+

$1/3$

+

$1/3$

=



Registration and atlasing require specific operations

Interpolation

Averaging

Smoothing

$1/3$

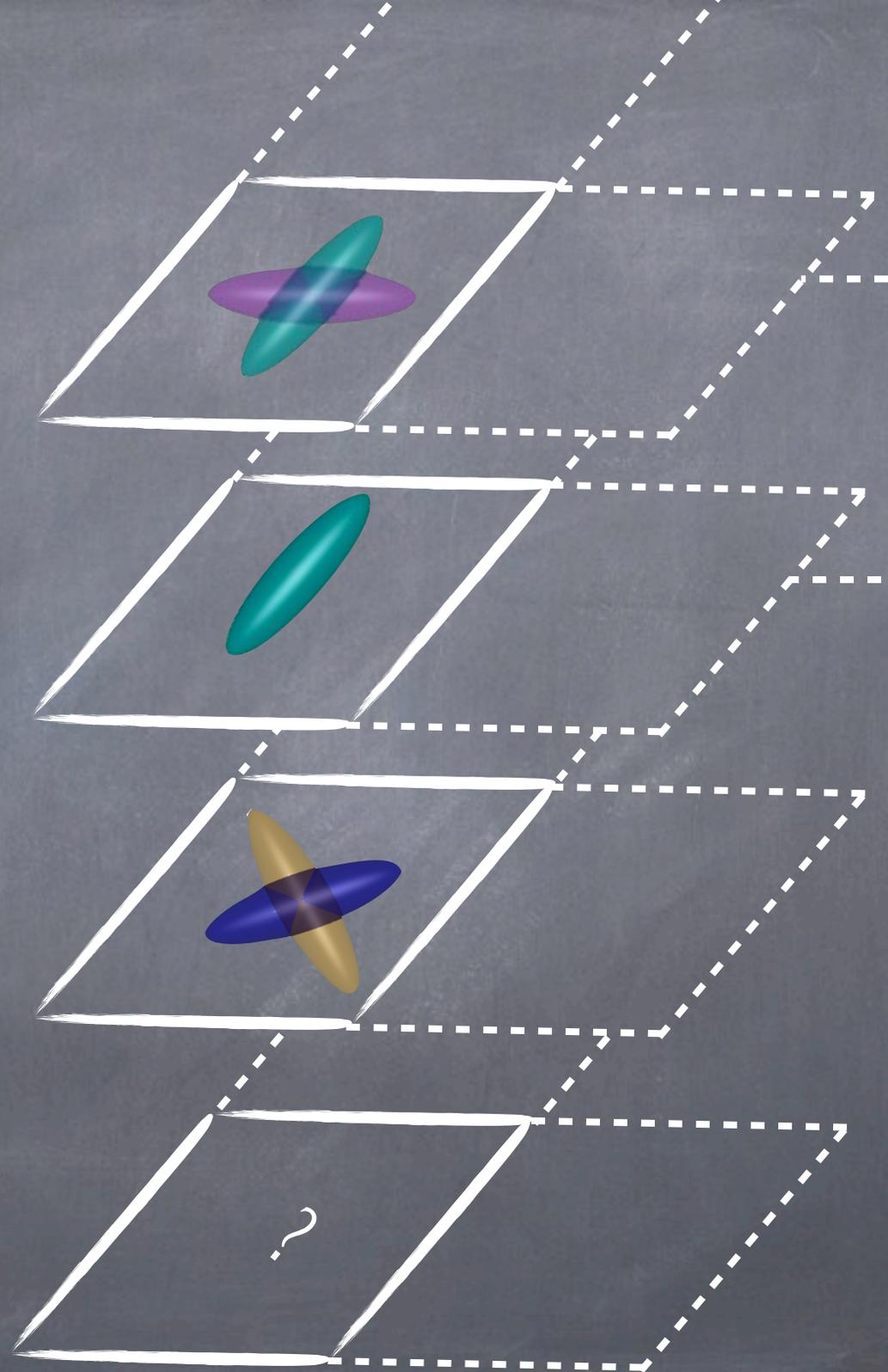
+

$1/3$

+

$1/3$

=



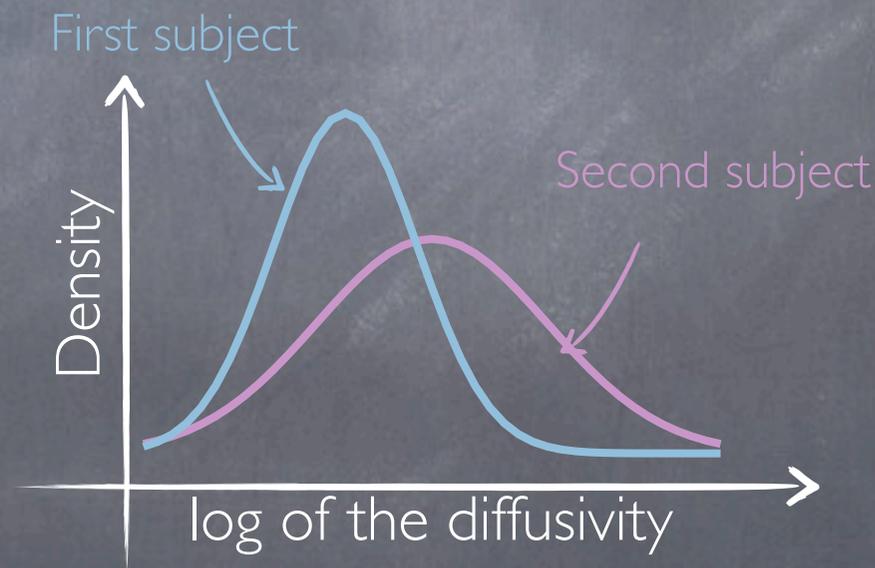
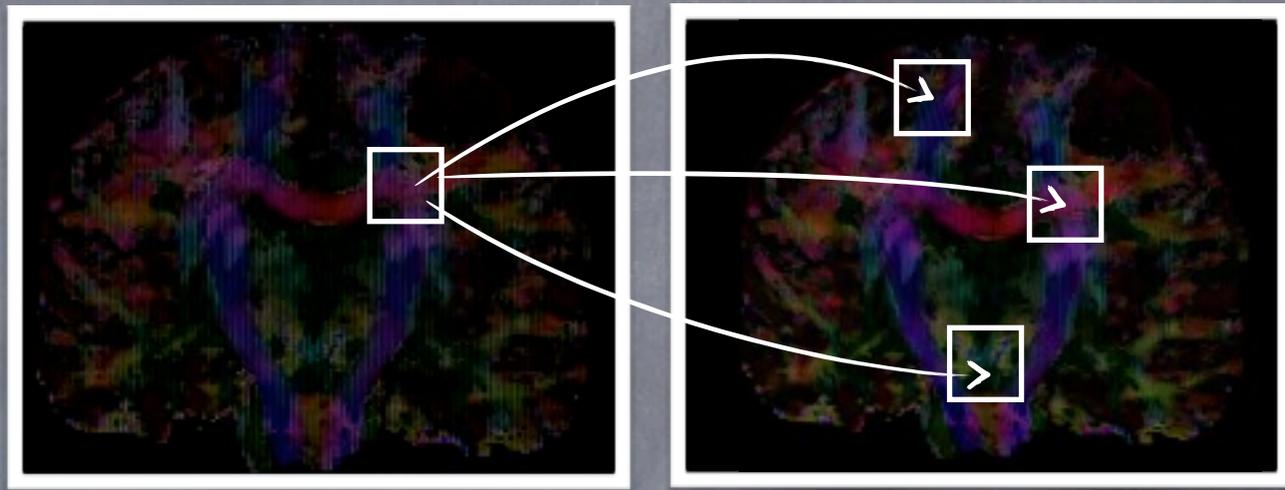
Registration and atlasing require specific operations

Interpolation

Averaging

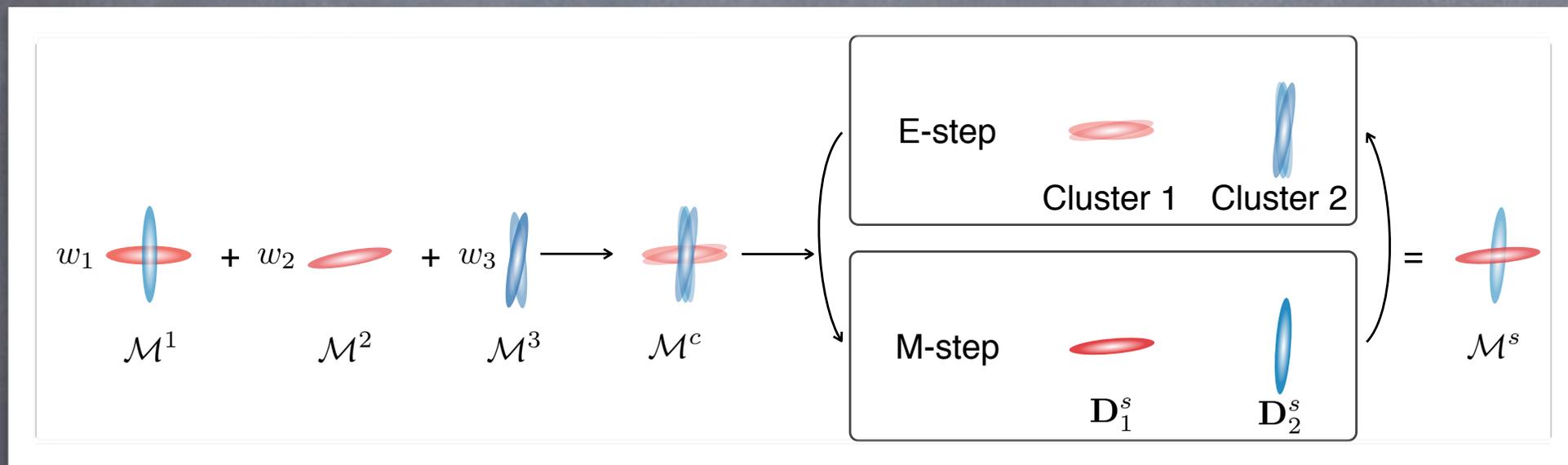
Smoothing

Similarity metric



Diffusivities differ between subjects

We proposed a mathematical framework that includes image analysis operators for multi-fascicle models



Minimization of the cumulative differential relative entropy

We proposed a mathematical framework that includes image analysis operators for multi-fascicle models

Generalized correlation coefficient

$$\rho(R, S) = \left\langle \frac{R - \mu_R}{\|R - \mu_R\|}, \frac{S - \mu_S}{\|S - \mu_S\|} \right\rangle$$

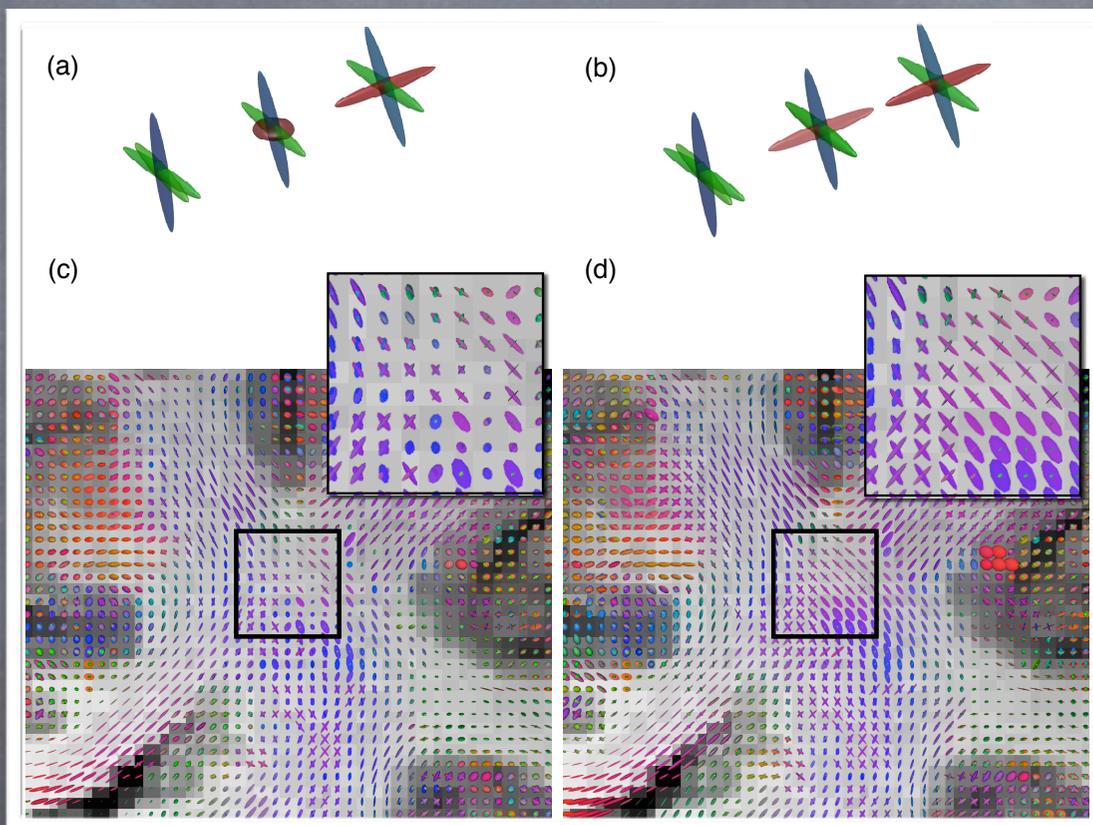
$$\rho(R, S) = m \left(\frac{R - m(R, T)T}{n_m(R - m(R, T)T)}, \frac{S - m(S, T)T}{n_m(S - m(S, T)T)} \right)$$

$$m(R, S) = \sum_{x \in \Omega} \max_{\pi} \sum_{i=1}^N f_i g_{\pi(i)} \left\langle \log \mathbf{D}_i^R, \log \mathbf{D}_{\pi(i)}^S \right\rangle$$

invariant under linear transformations of the log-eigenvalues

Our framework enables accurate registration and atlas construction from multi-fascicle models

Experiment I: Interpolation Error

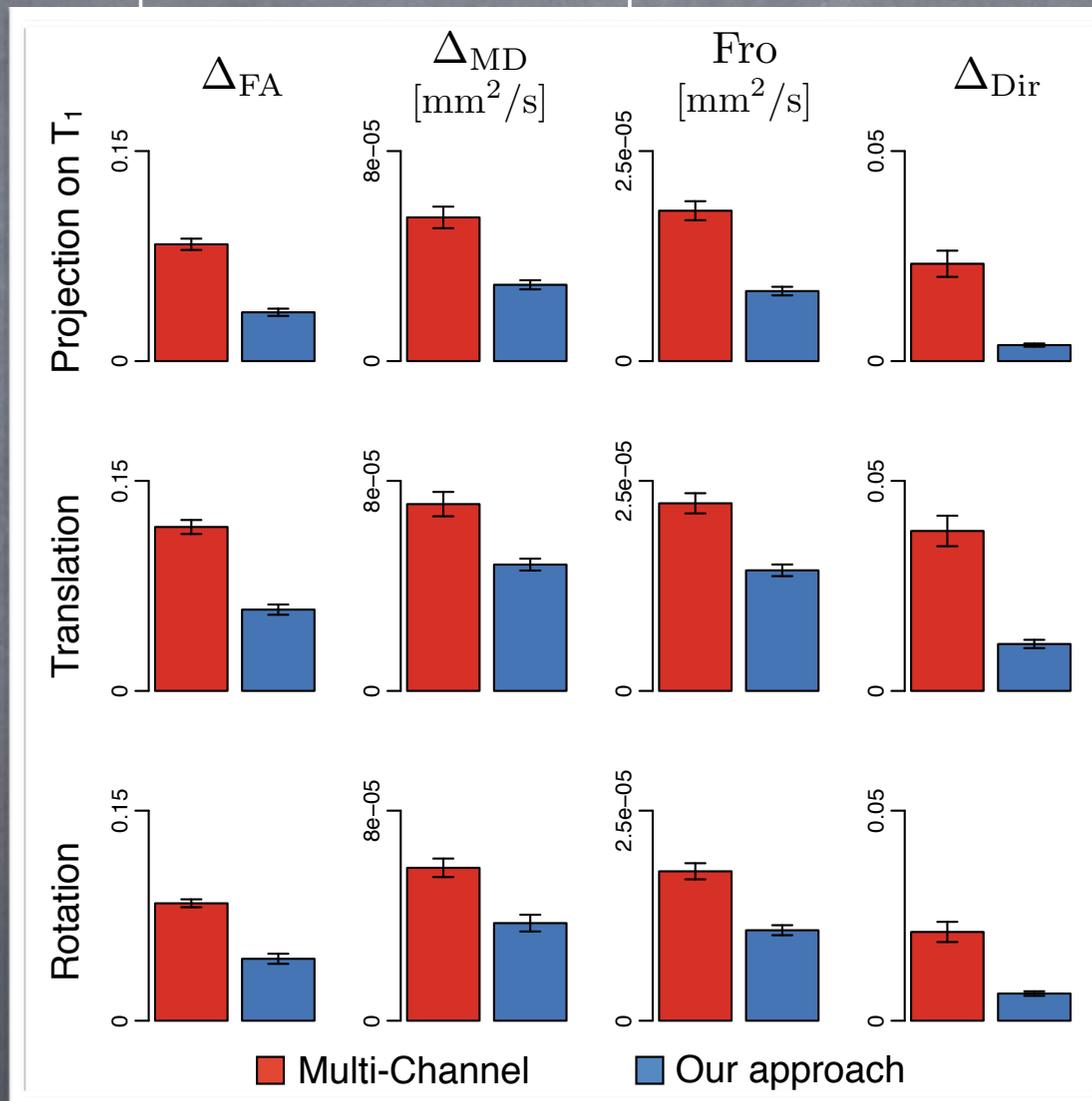


Multi-channel

Our approach

Our framework enables accurate registration and atlas construction from multi-fascicle models

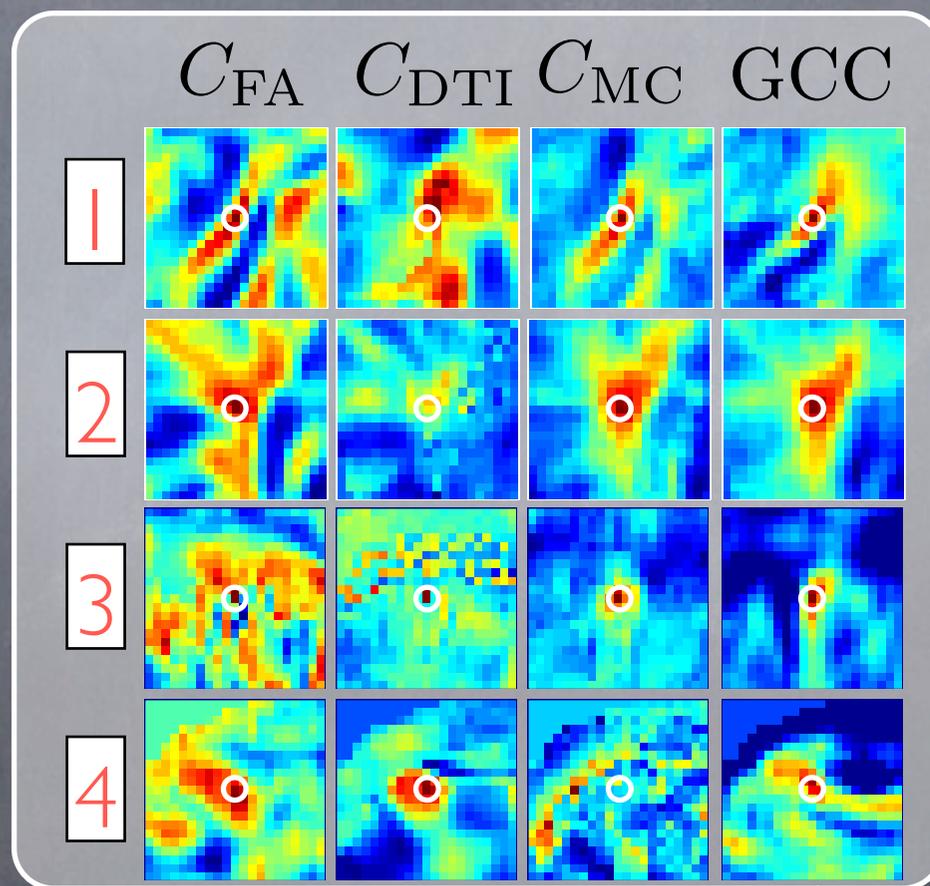
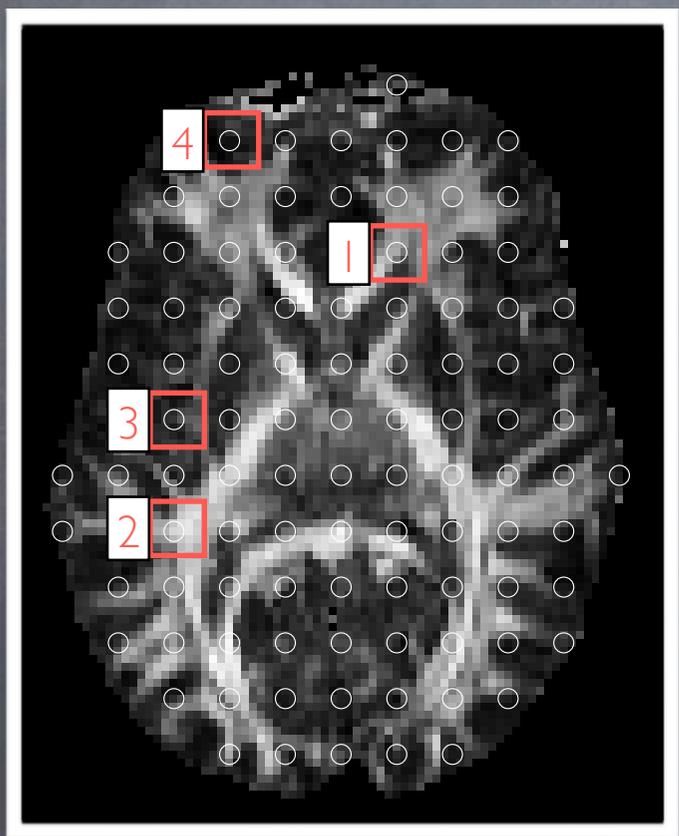
Experiment I: Interpolation Error



Our framework enables accurate registration and atlas construction from multi-fascicle models

Experiment 2: Scan-Rescan to test the similarity metric

Saliency maps

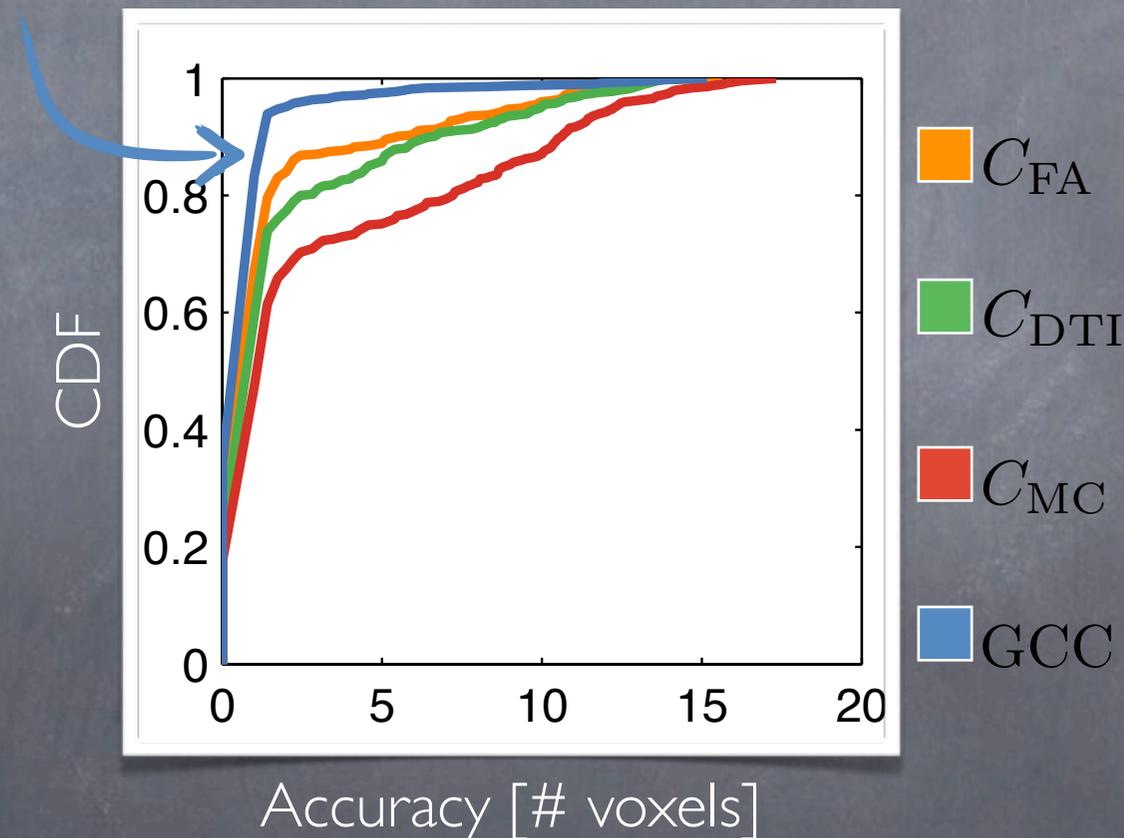


Our metric

Our framework enables accurate registration and atlas construction from multi-fascicle models

Experiment 2: Scan-Rescan to test the similarity metric

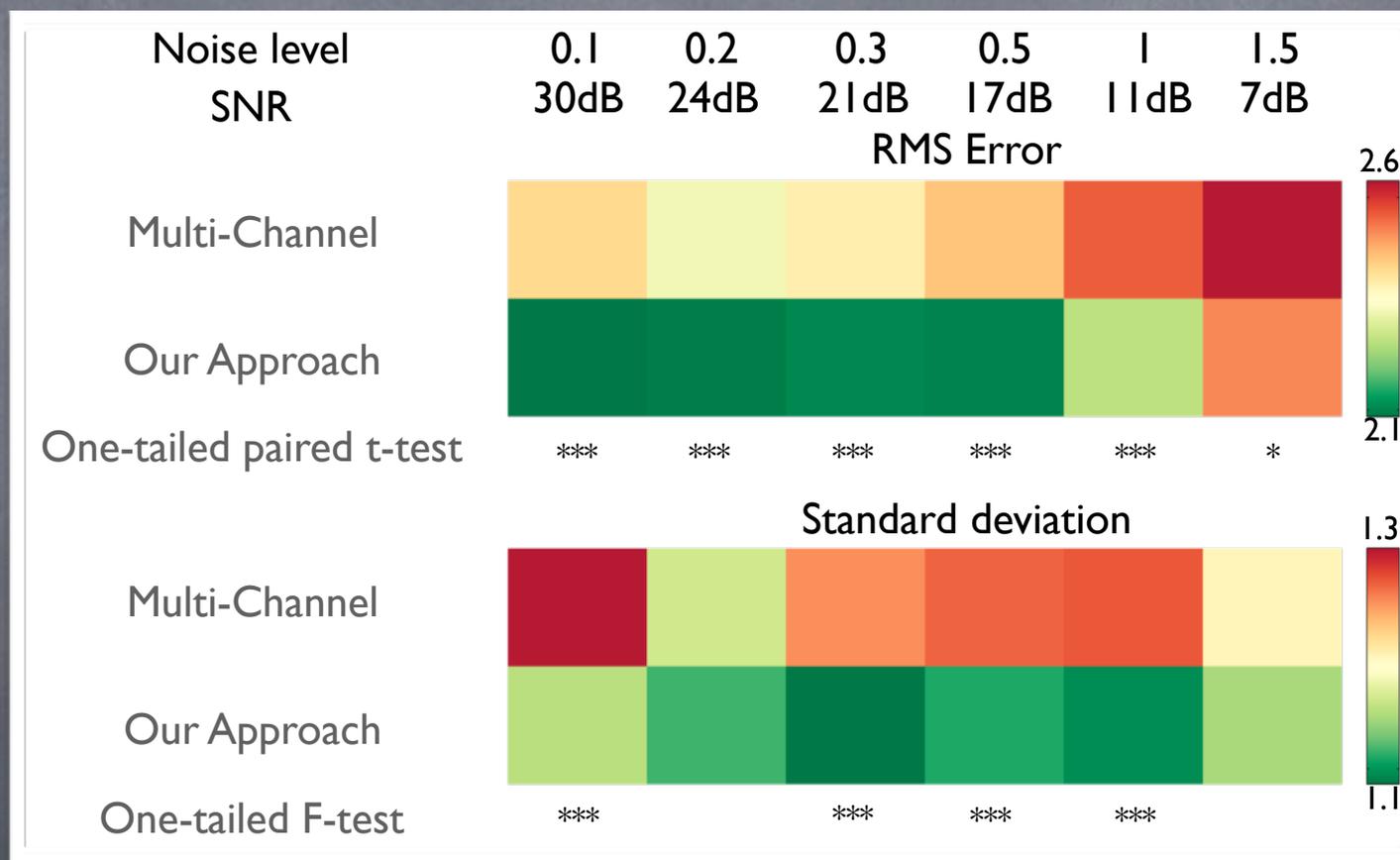
Our metric leads to significantly smaller registration errors



Our framework enables accurate registration and atlas construction from multi-fascicle models

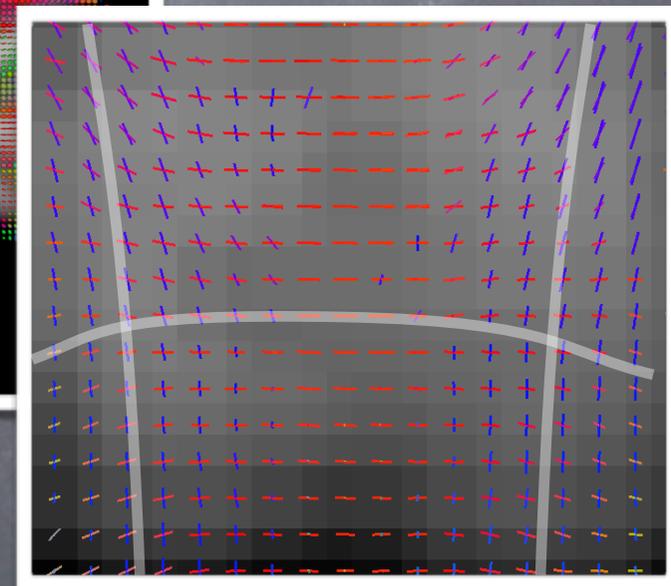
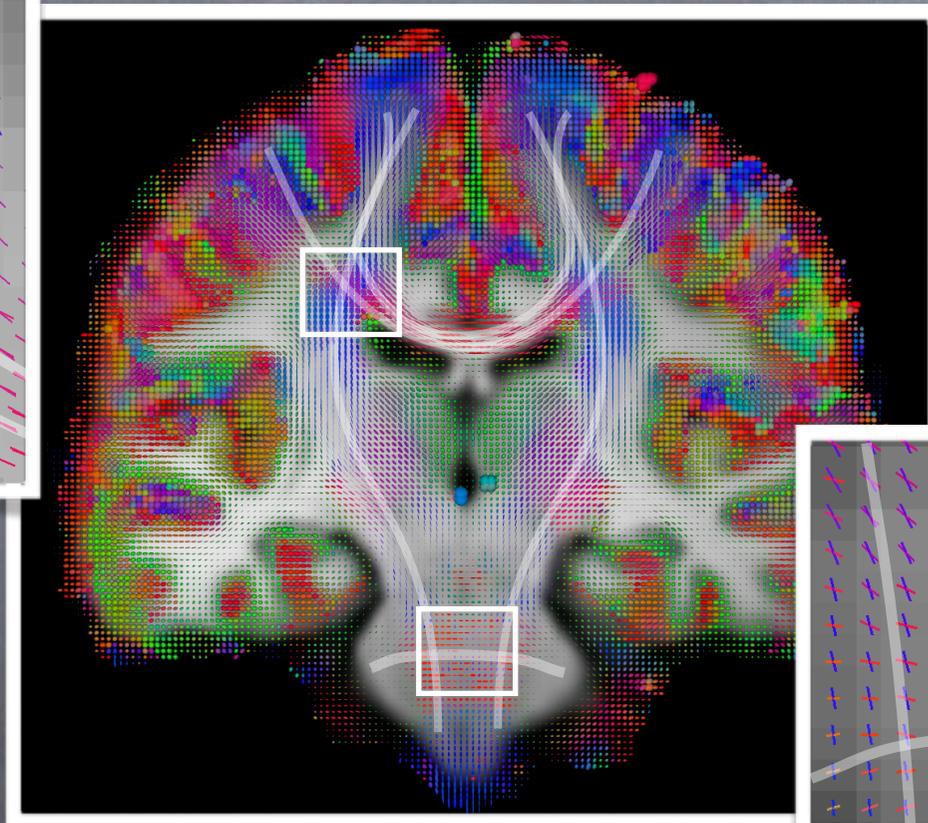
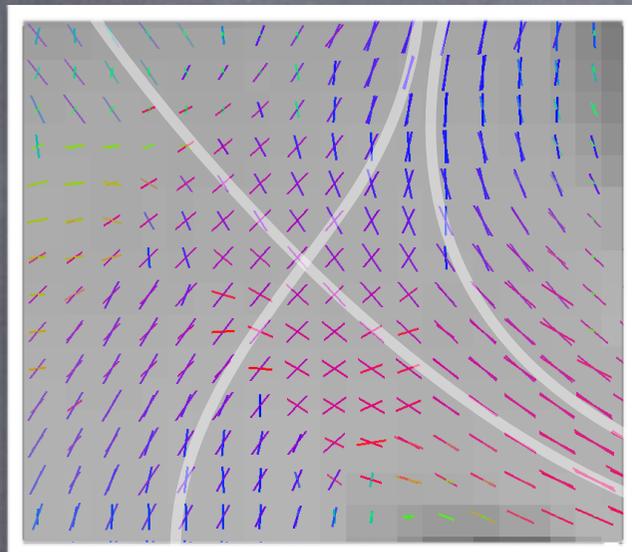
Experiment 3: 1,440 synthetic field registrations

Our method performs significantly better



Our framework enables accurate registration and atlas construction from multi-fascicle models

Our framework enables the construction of a multi-fascicle atlas



Regions with known crossing fascicles are well represented in the atlas

New assets and capabilities to conduct population studies of the brain microstructure

1. Selection of the appropriate model
2. Registration and atlas construction
- 3. Statistical analysis of microstructure**
4. Estimation from single b-value data

Features of the brain microstructure may relate to a particular fascicle or to the surrounding volume

Grey matter

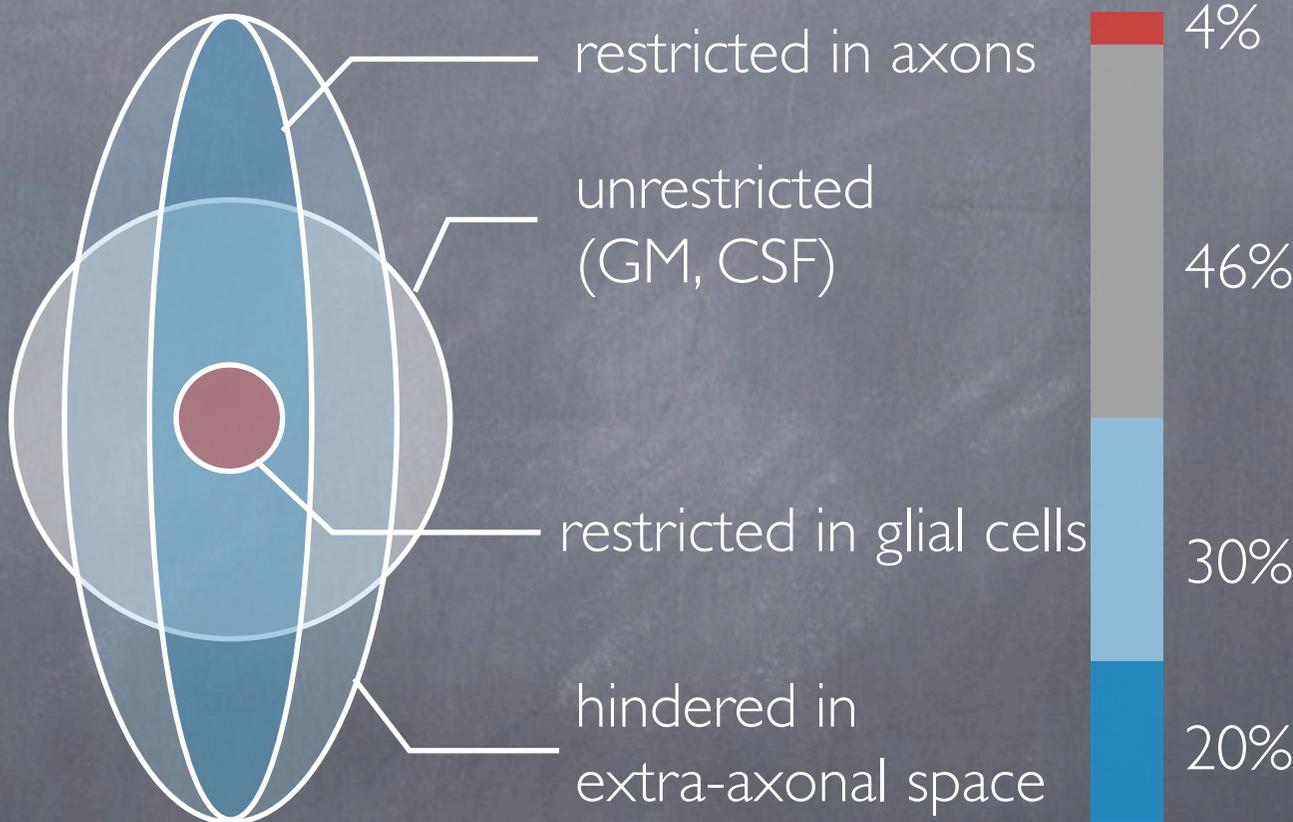
Astrocyte

Axon

Myelin

Oligodendrocyte

Microglia



We proposed a system of two methods for the statistical analysis of microstructural features

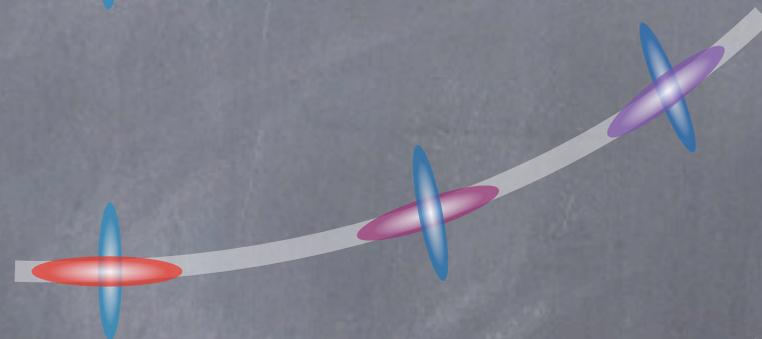
Fascicle-based spatial statistics

for properties related to fascicles

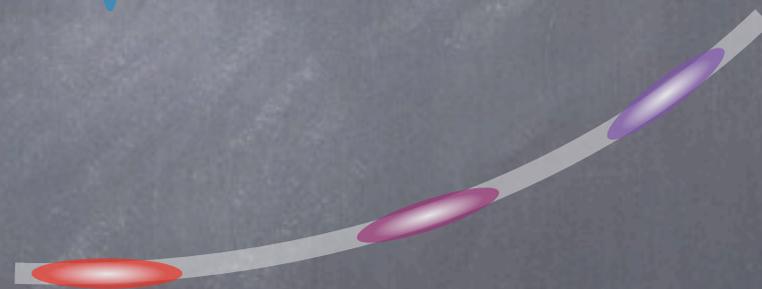
1. tractography



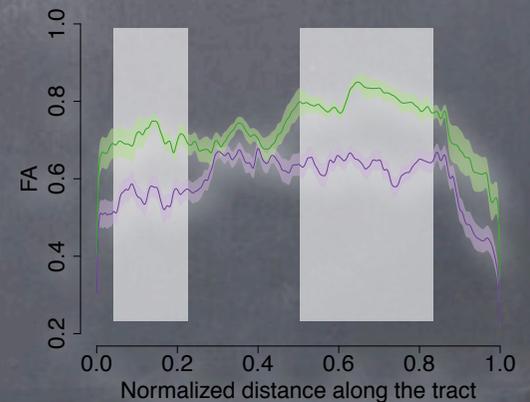
2. interpolation



3. selection



4. cluster-based statistics



We proposed a system of two methods for the statistical analysis of microstructural features

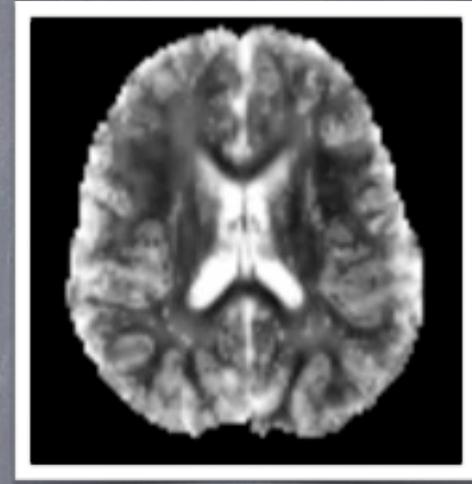
Isotropic diffusion analysis

for properties related to volume

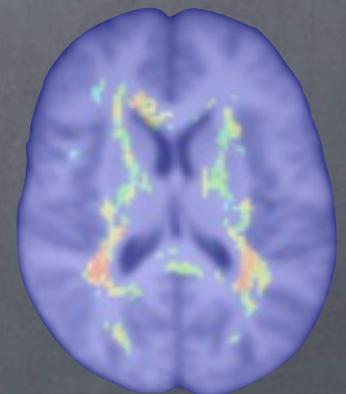
1. Feature extraction

2. Feature transformation

3. cluster-based statistics

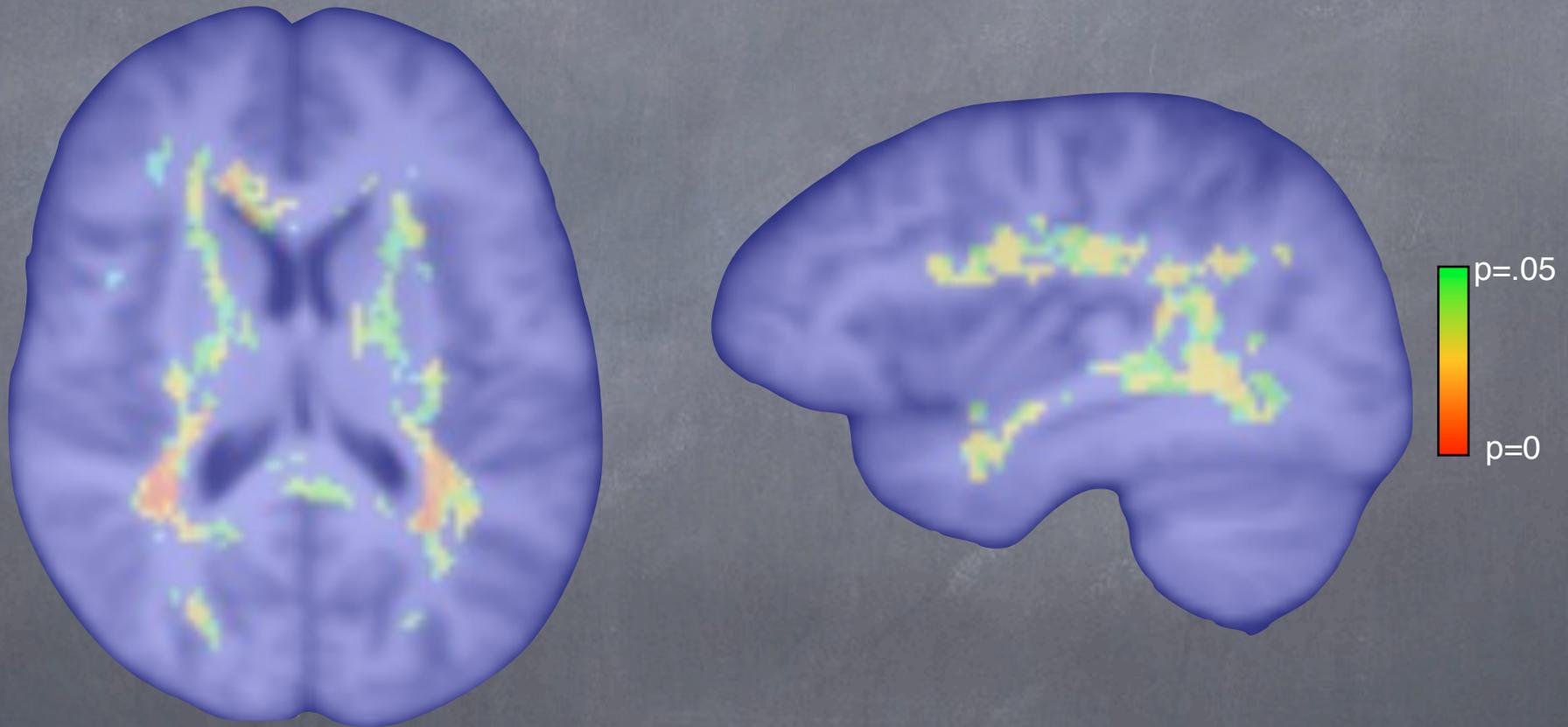


$$l_{\text{iso}} = \text{logit } f_{\text{iso}}$$



Our system detects differences between healthy controls and children with autism

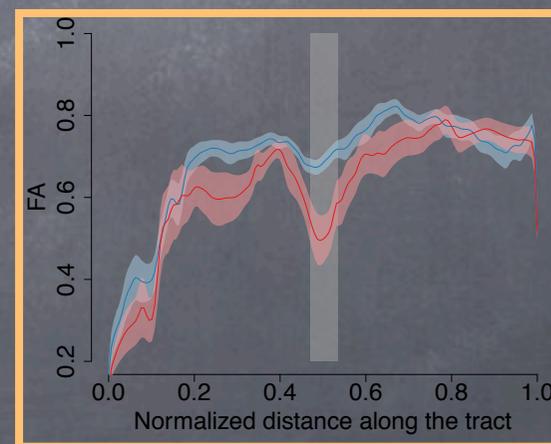
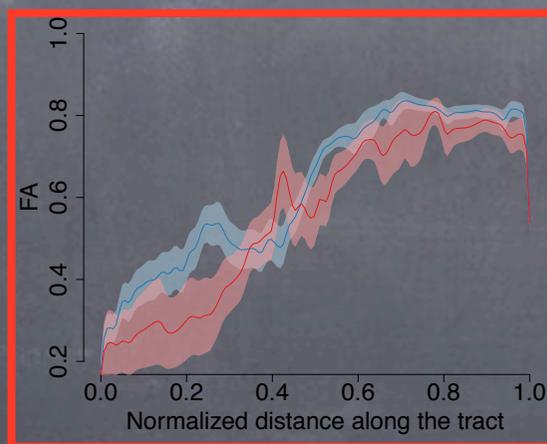
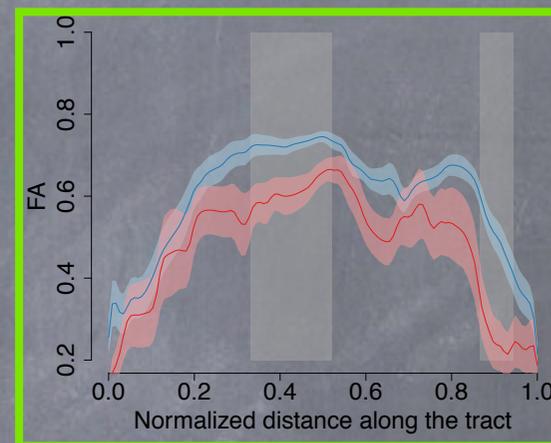
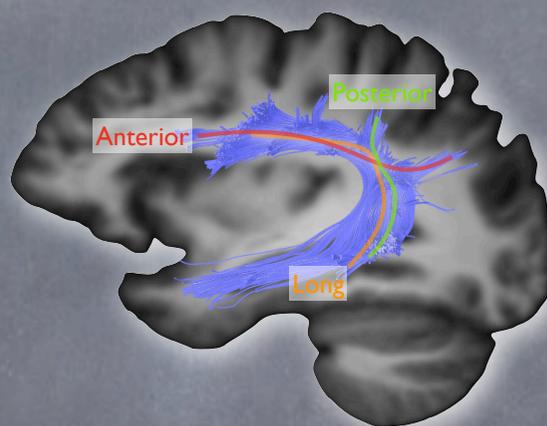
IDA reveals regions with abnormally high fraction of molecules diffusing freely, mostly in the dorsal language pathway.



This finding may point to the hypothesis that autism results from an autoimmune response.

Our system detects differences between healthy controls and children with autism

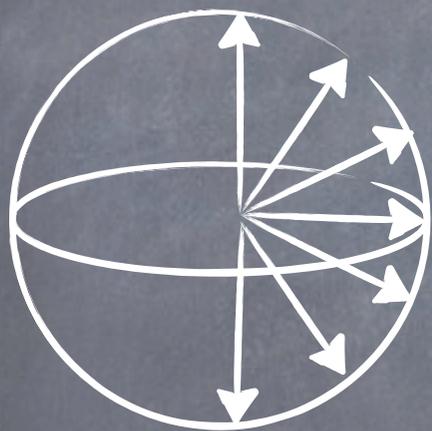
FBSS reveals few clusters of significantly lower fractional anisotropy along the dorsal language pathway.



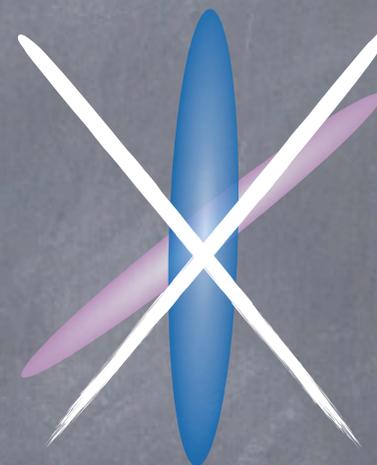
New assets and capabilities to conduct population studies of the brain microstructure

1. Selection of the appropriate model
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4. Estimation from single b-value data

Multi-fascicle models cannot be directly estimated from data at a single b-value

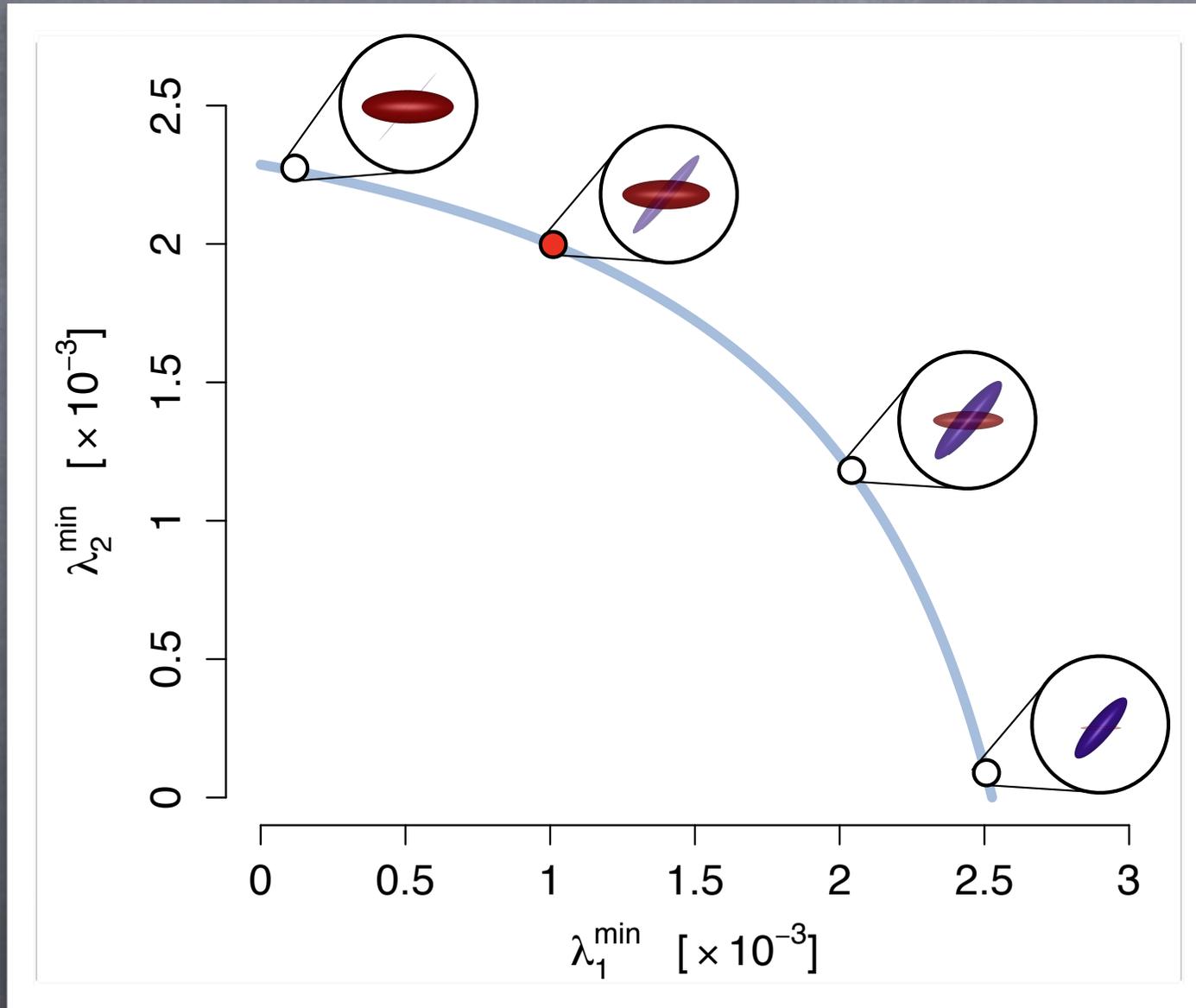


Single gradient shell

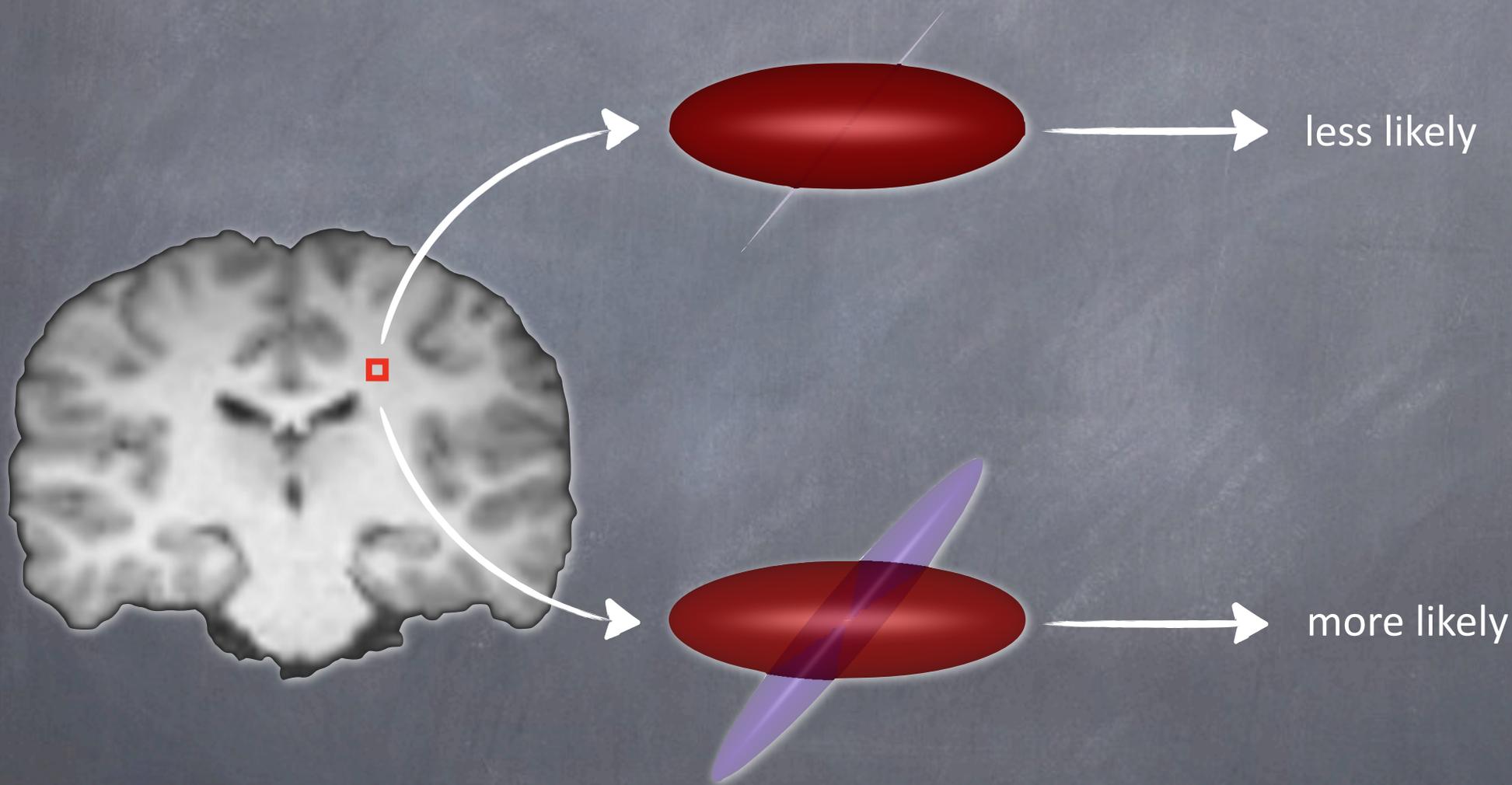


The problem is ill-posed

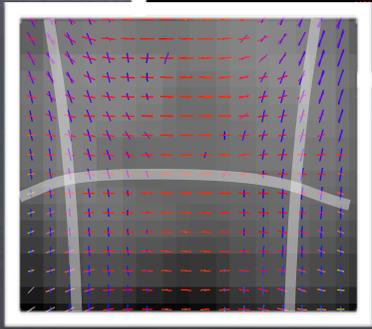
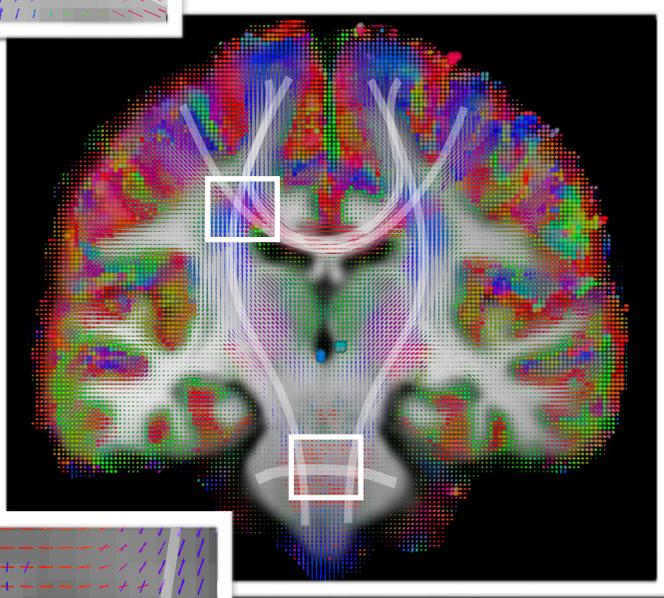
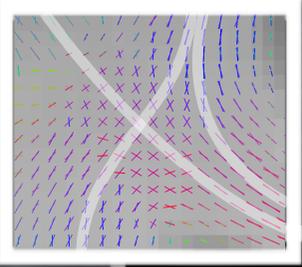
All those models produce the same signal



Not all models are as likely given the known anatomy



We proposed to incorporate prior information from other subjects scanned at multiple b-values



Each voxel of the atlas contains a parametric distribution of the parameters

We proposed to incorporate prior information from other subjects scanned at multiple b-values

Each voxel of the atlas contains a parametric distribution of the parameters

$$p(\mathbf{f}, \mathbf{D} | \text{Data}) \propto p(\text{Data} | \mathbf{f}, \mathbf{D}) p(\mathbf{f}, \mathbf{D})$$

Likelihood \nearrow \nwarrow Prior

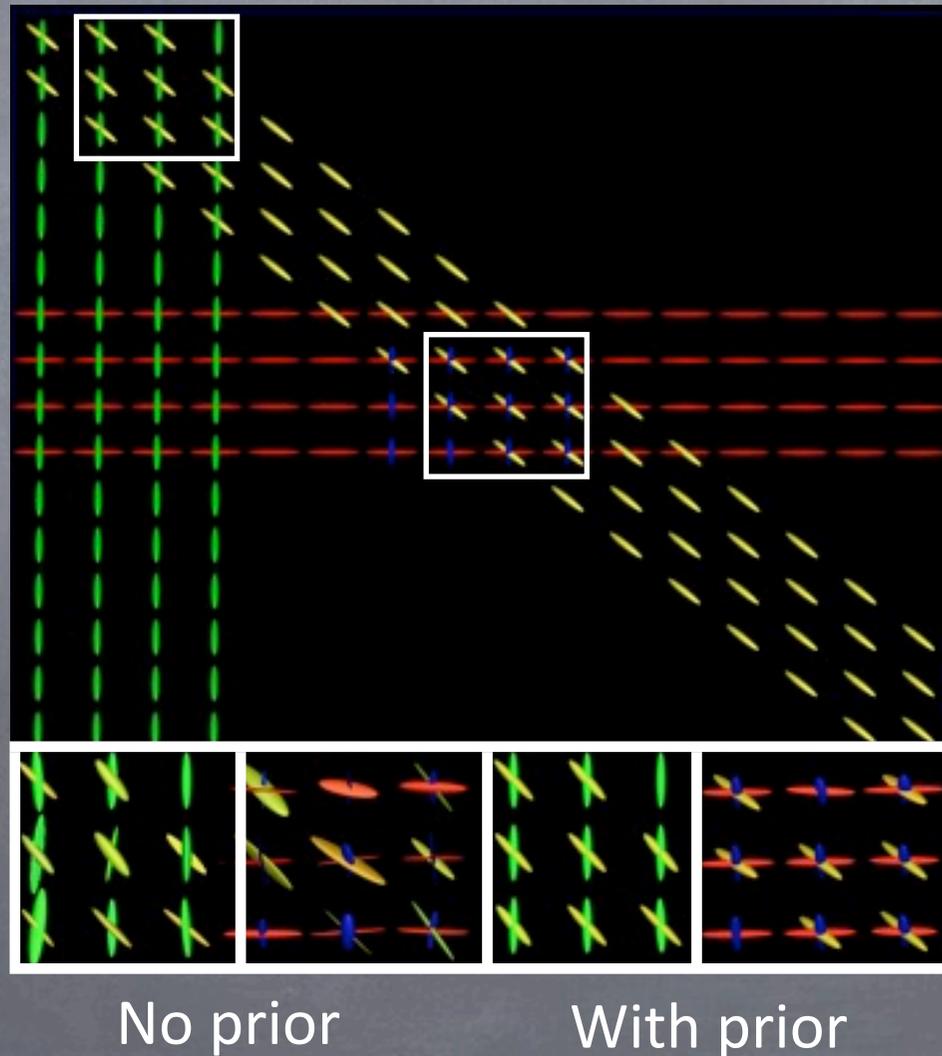
$p(\mathbf{f}, \mathbf{D})$ is informed by the atlas

$$p(\mathbf{f}, \mathbf{D}) = p(\mathbf{f}) p(\mathbf{D})$$

Dirichlet \nearrow \nwarrow Matrix-variate Gaussian

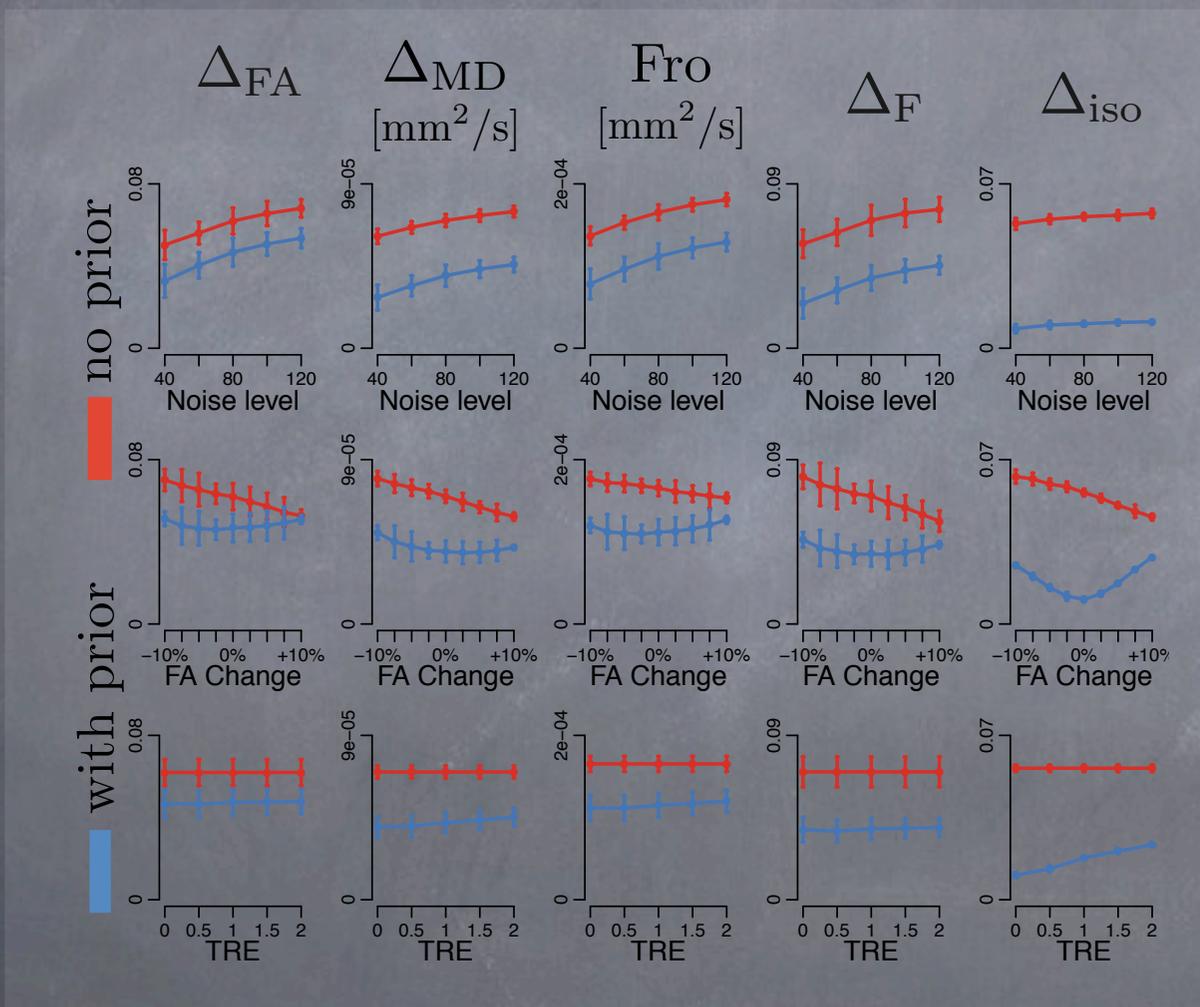
The population-informed prior significantly improves the accuracy of the estimation

Experiment I: Synthetic data



The population-informed prior significantly improves the accuracy of the estimation

Experiment I: Synthetic data



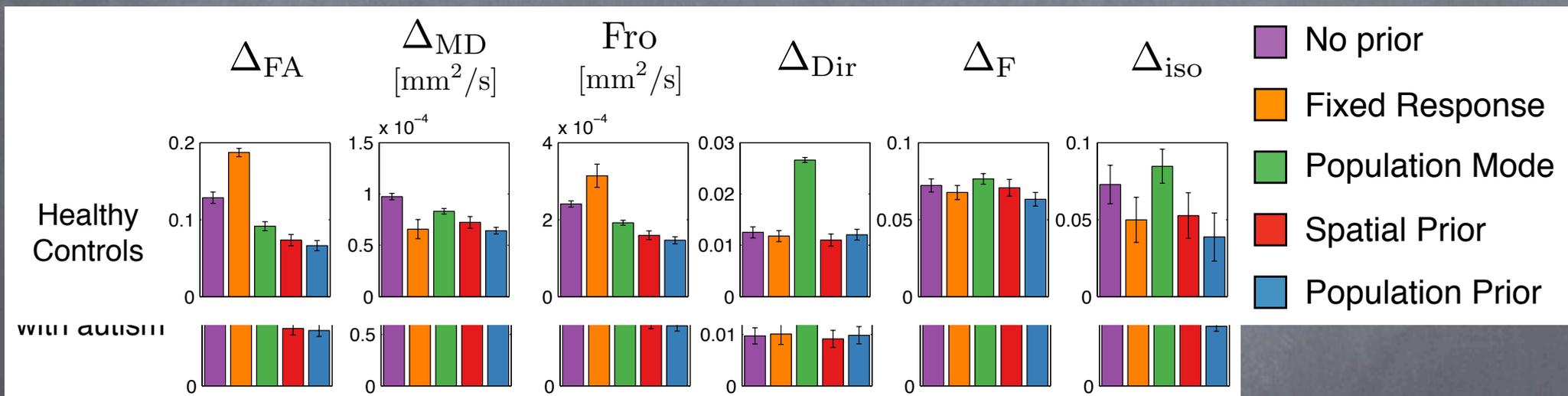
Under noise

Under group differences

Under registration error

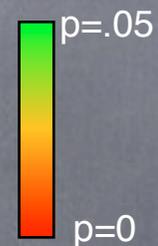
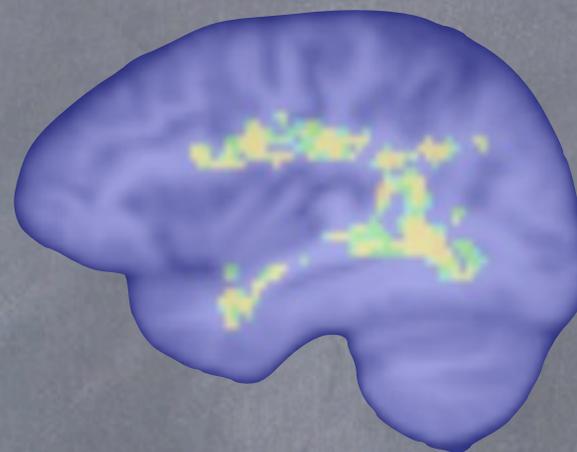
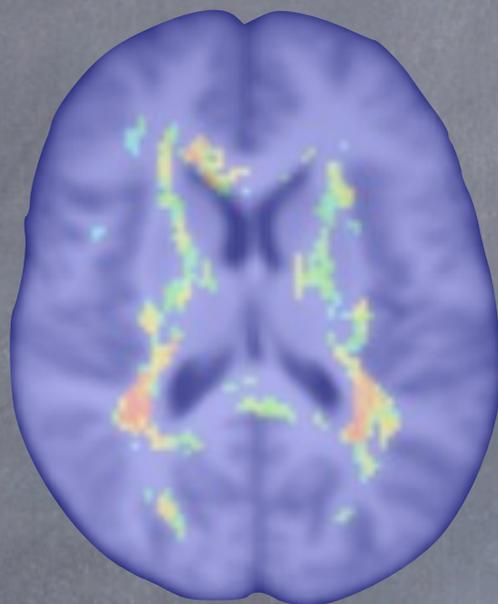
The population-informed prior significantly improves the accuracy of the estimation

Experiment 2: In-vivo data

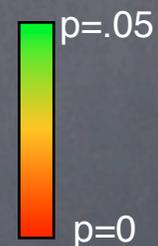
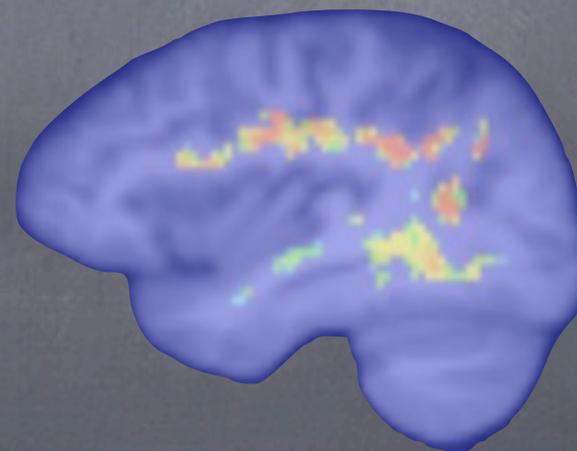
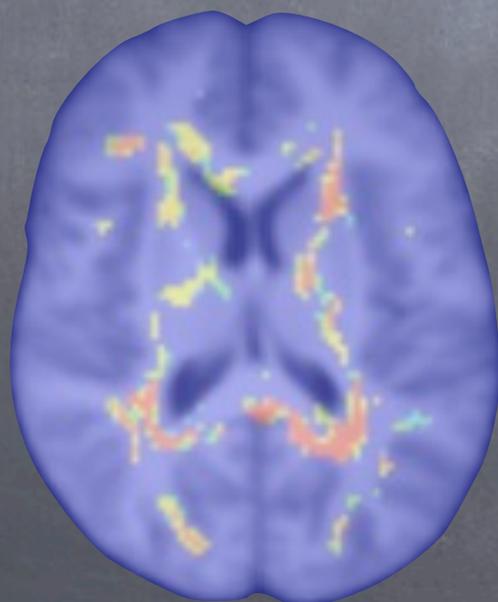


Detected group differences are remarkably close to those found with multiple b-values

With multiple
b-values

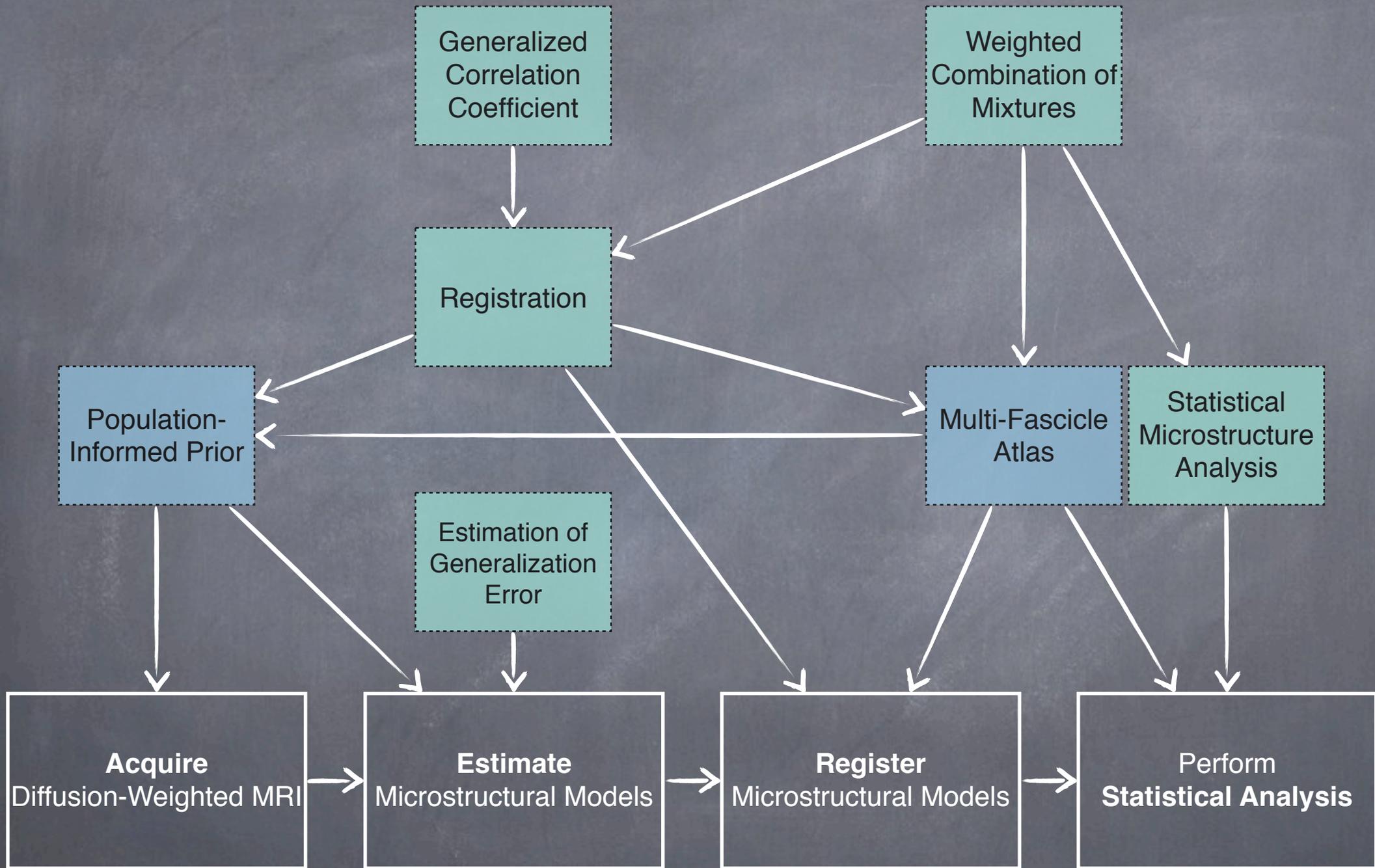


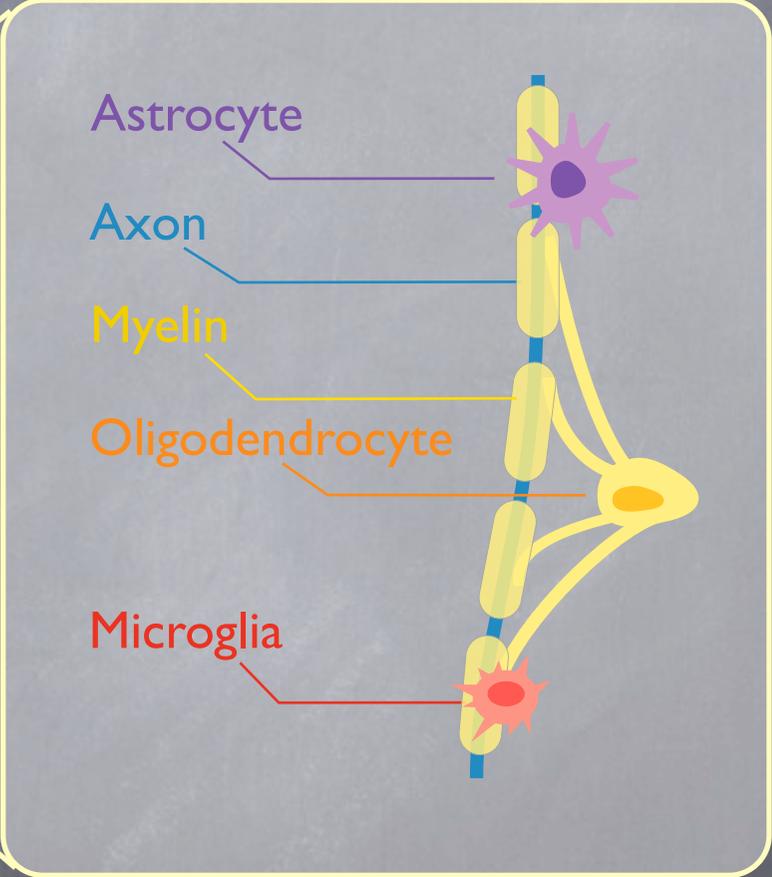
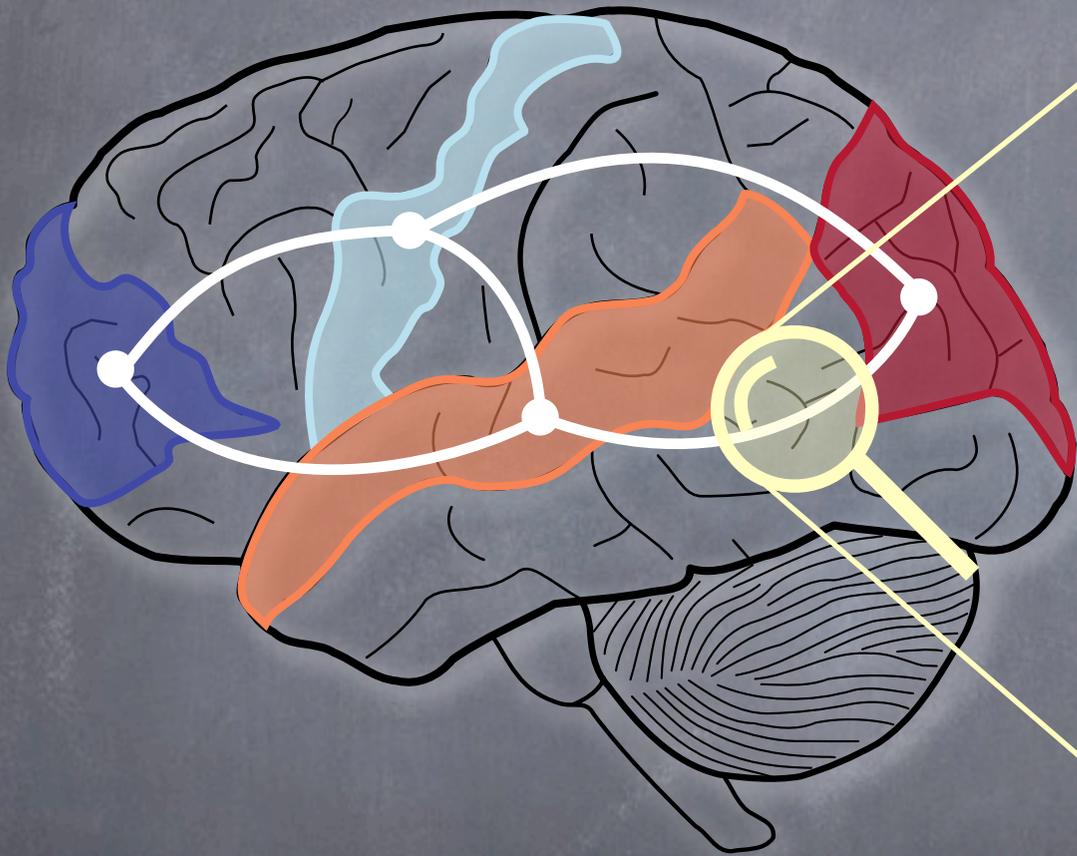
With a single
b-value



New assets and capabilities to conduct population studies of the brain microstructure

1. Selection of the appropriate model
2. Registration and atlas construction
3. Statistical analysis of microstructure
4. Estimation from single b-value data





Thank you

Journals

- Taquet et al., Neuroimage (under review)
- Taquet et al., IEEE TMI, 2013
- Peters*, Taquet* et al., Future Neurology, 2013
- Taquet and Peters, Médecine/Sciences, 2013
- Peters*, Taquet*, et al., BMC Medicine, 2013

Peer-reviewed conference abstracts

- Taquet et al., OHBM 2013
- Taquet et al., ISMRM 2013
- Scherrer et al., ISMRM 2013
- Peters*, Taquet* et al., American Acad. of Neurology 2013
- Peters*, Taquet* et al. Child Neurology Society, 2012
- Taquet et al., CAOS 2010

Full-length peer-reviewed conference papers

- Taquet et al., MICCAI 2013
- Scherrer et al., MICCAI 2013
- Scherrer*, Taquet* et al., IPMI 2013
- Taquet et al., MICCAI 2012
- Taquet et al., IEEE ISBI 2012
- Taquet et al., IEEE MMBIA 2012
- Taquet et al., MICCAI 2011
- Taquet et al., IEEE ICIP 2011
- Taquet et al., CTIC 2010
- Taquet, IEEE Melecon 2010

* Equal contributions

