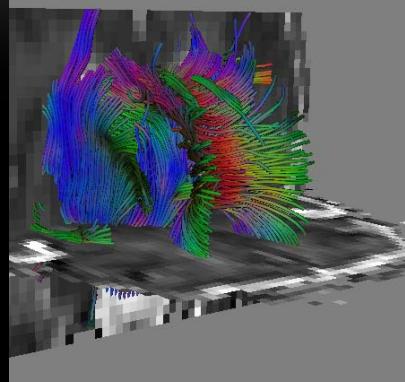
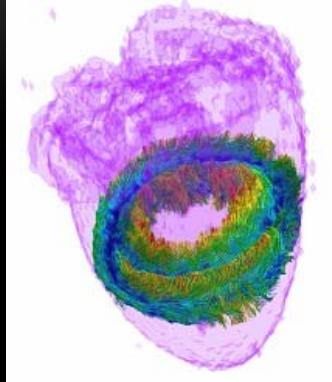
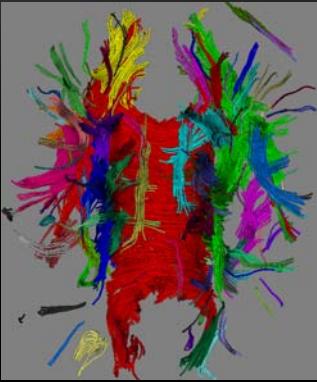


Diffusion Tensor Imaging Visualization Techniques and Applications



Tim Peeters (t.peeters@tue.nl) - **Anna Vilanova** (a.vilanova@tue.nl)

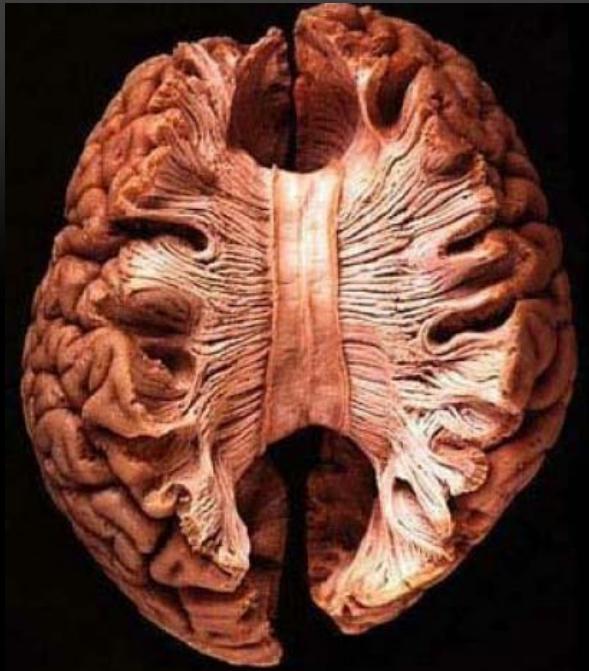
BioMedical Image Analysis (bmia.bmt.tue.nl)

Eindhoven University of Technology, The Netherlands

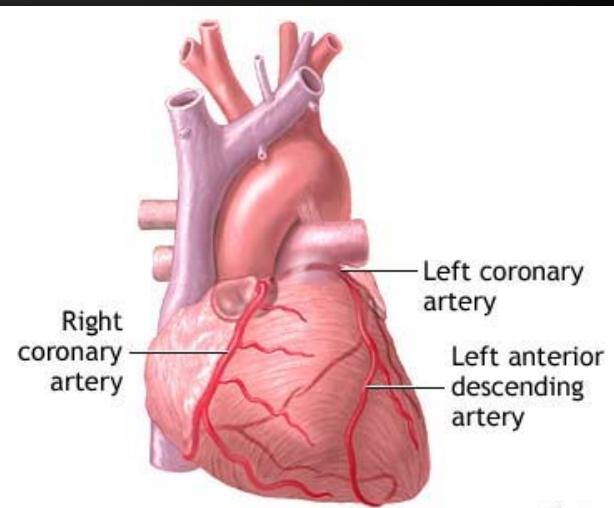
Overview

- Diffusion Tensor Imaging (DTI) data
- DTI visualization techniques
- Applications: newborn and ischemic heart
- Fiber clustering
- DTI segmentation

Motivation



T.H. Williams et al.



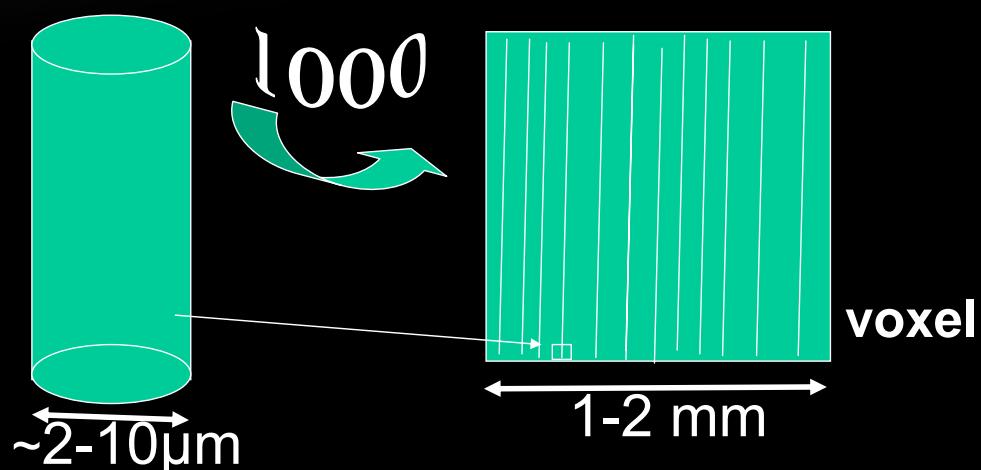
ADAM

<http://www.shands.org/>

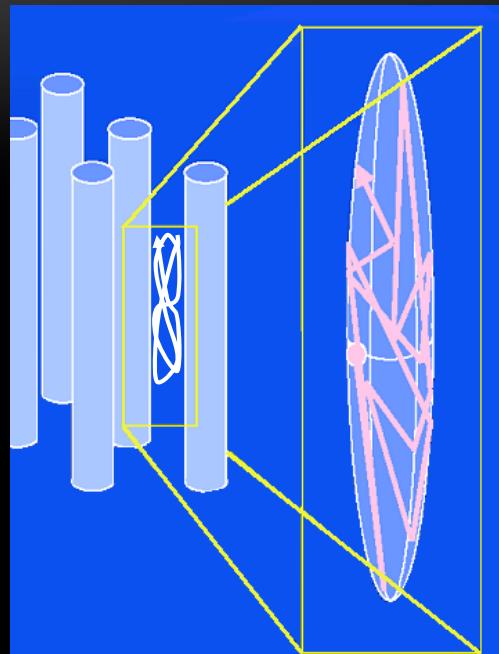
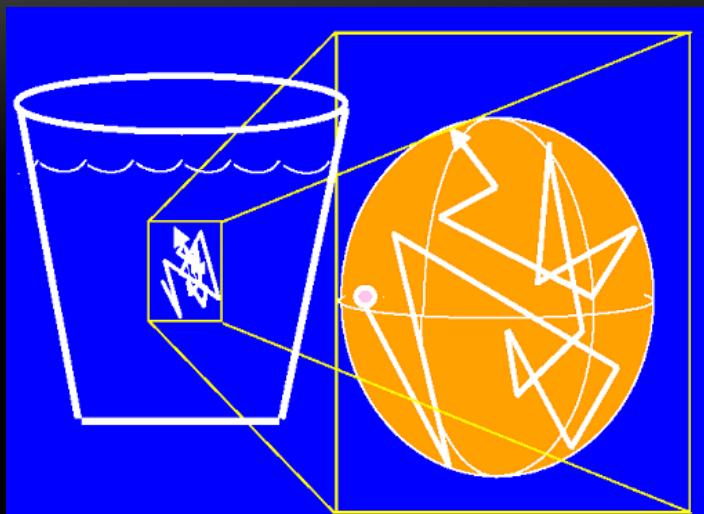
Motivation MRI and Diffusion Tensor Imaging

Fibers – Micrometers ($\sim 2\text{-}10\mu\text{m}$)

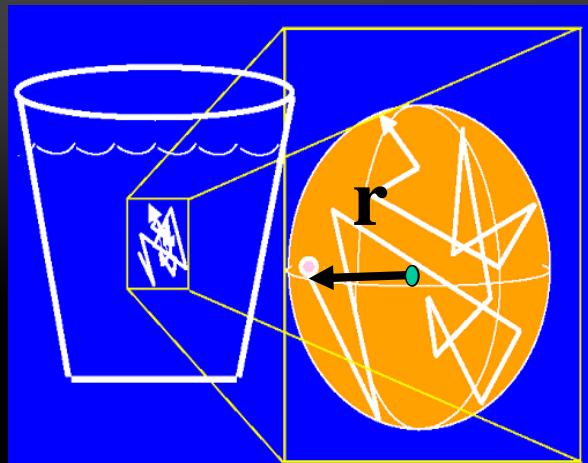
Scanner (MR) – Millimeters ($\sim 1\text{-}2\text{ mm}$)



Water Diffusion Brownian Motion



Fick's Law



Solution -3D Gaussian

$$P(\mathbf{r}, t) = \frac{1}{(4\pi D t)^{3/2}} e^{-\frac{1}{4t} \mathbf{r}^2}$$

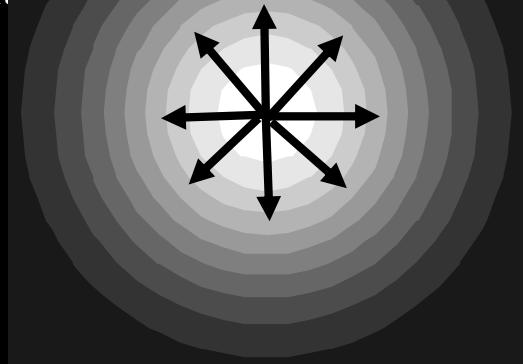
$$\frac{\partial}{\partial t} P(\mathbf{r}, t) = D \cdot \nabla^2 P(\mathbf{r}, t)$$

t Diffusion time

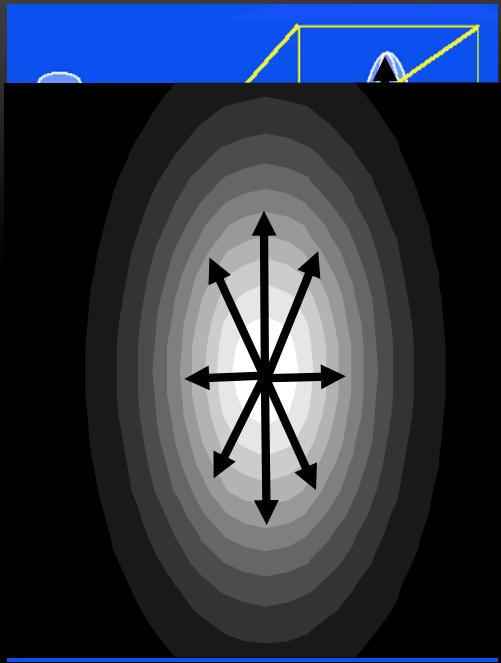
D Diffusion coefficient (mm^2/s)

P Probability density function

transient



Anisotropic Diffusion

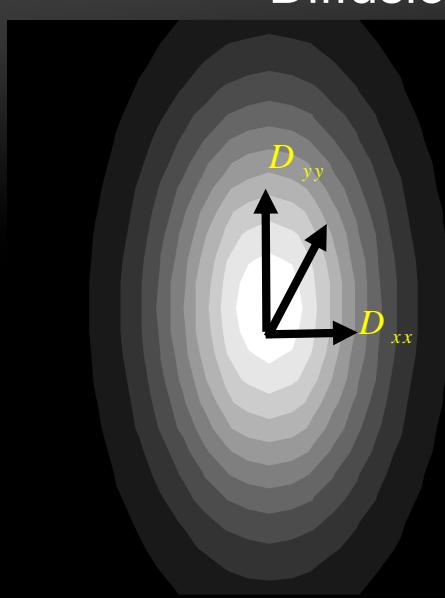


$$P(\mathbf{r}, t) = \frac{1}{(4\pi D t)^{3/2}} e^{-\frac{1}{4t} \mathbf{r}^2 D^{-1}}$$

$$P_i(\mathbf{r}_i, t) = \frac{1}{(4\pi D_i t)^{3/2}} e^{-\frac{1}{4t} \mathbf{r}_i^2 D_i^{-1}}$$

Indicates the distance squared of the vector

Anisotropic Diffusion – Diffusion Tensor



Diffusion Tensor

$$\mathbf{D} = \begin{bmatrix} D_{xx} & D_{xy} & D_{xz} \\ D_{yx} & D_{yy} & D_{yz} \\ D_{zx} & D_{zy} & D_{zz} \end{bmatrix}$$

$$D_i = \mathbf{r}_i^t \mathbf{D} \mathbf{r}_i$$

$$P(\mathbf{r}, t) = \frac{1}{(4\pi |\mathbf{D}| t)^{3/2}} e^{-\frac{1}{4t} \mathbf{r}^t \mathbf{D}^{-1} \mathbf{r}}$$

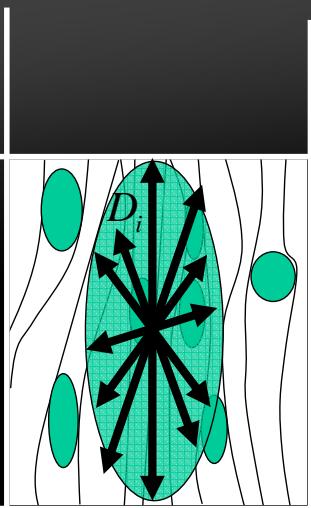
- Measure Diffusion Weighted signal S_i in a given direction
- Stejskal-Tanner relationship attenuation signal S_i to diffusion coefficient D_i

$$S_i = S_0 e^{-bD_i} \quad \text{where } S_0 \text{ not diffusion weighted value}$$

b protocol parameter (diffusion time, ...)

is often called ADC_i (Apparent Diffusion Coefficient) – D_i
diffusion in a given direction

MRI-Diffusion Measurement



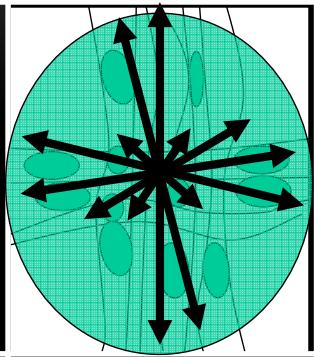
Axis indicates
preferred direction

Measure ADC_i in a lot of directions (Minimum 6)

Fit **D**
2nd Order Tensor
Symmetric
Positive Definite

Assume Gaussian within a voxel

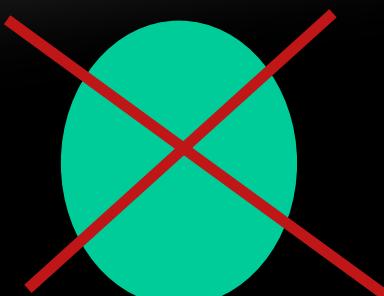
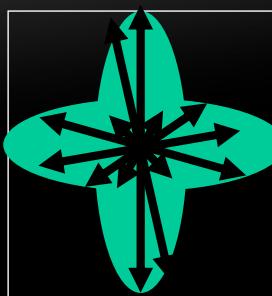
What problems does the Gaussian model have?



No preferred diffusion direction!

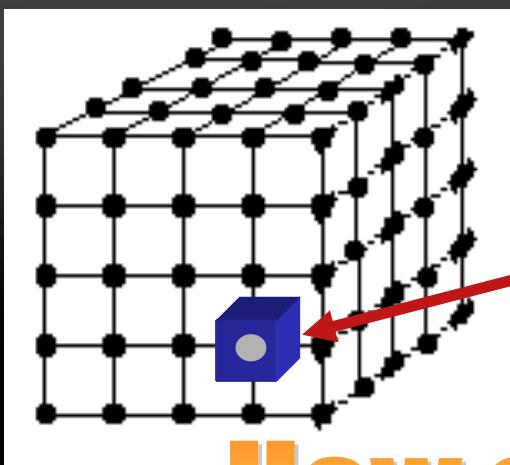
Other models

- HARDI - use other models for the probability density function.



- We will just talk about the Gaussian model!

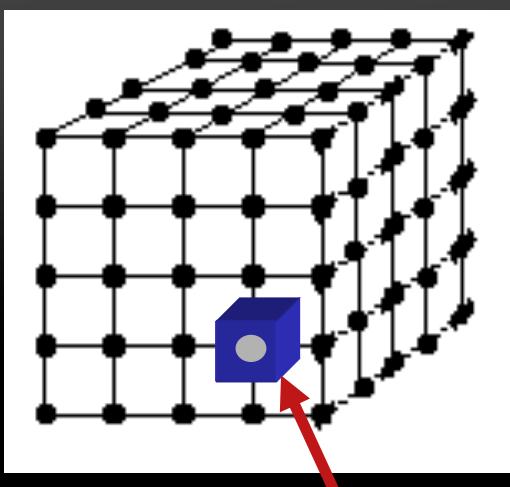
Diffusion Tensor Imaging Visualization



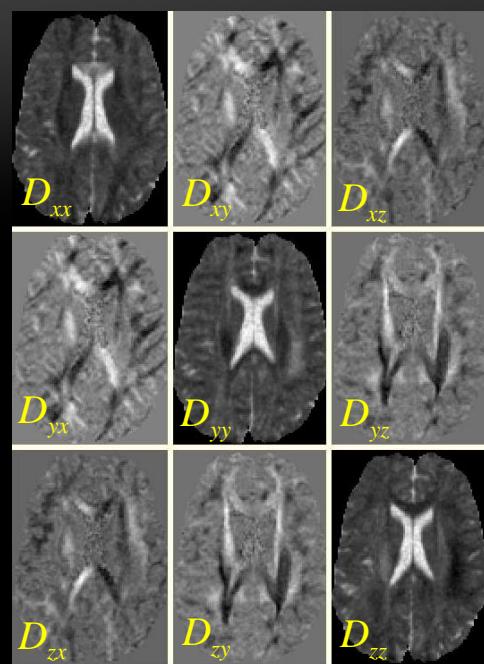
$$\mathbf{D} = \begin{bmatrix} D_{xx} & D_{xy} & D_{xz} \\ D_{yx} & D_{yy} & D_{yz} \\ D_{zx} & D_{zy} & D_{zz} \end{bmatrix}$$

**How are we
going to show this?**

Diffusion Tensor Imaging Visualization



$$\mathbf{D} = \begin{bmatrix} D_{xx} & D_{xy} & D_{xz} \\ D_{yx} & D_{yy} & D_{yz} \\ D_{zx} & D_{zy} & D_{zz} \end{bmatrix}$$



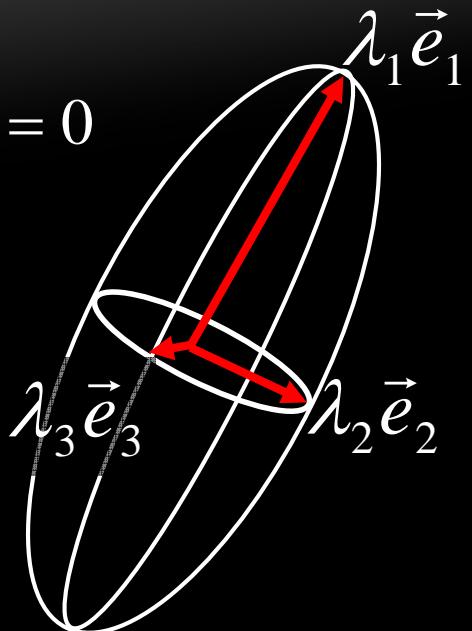
Eigenanalysis

$$\mathbf{D}\vec{e}_i = \lambda_i \vec{e}_i \quad \det(\lambda \mathbf{I} - \mathbf{D}) = 0$$

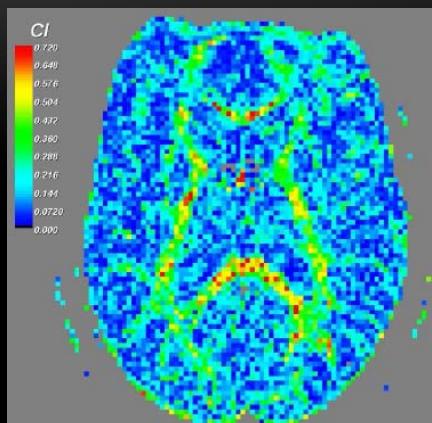
Eigenvalues

$$\lambda_1 \geq \lambda_2 \geq \lambda_3 \geq 0$$

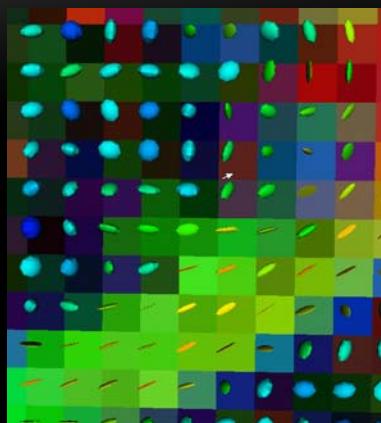
$$\vec{e}_1, \vec{e}_2, \vec{e}_3$$



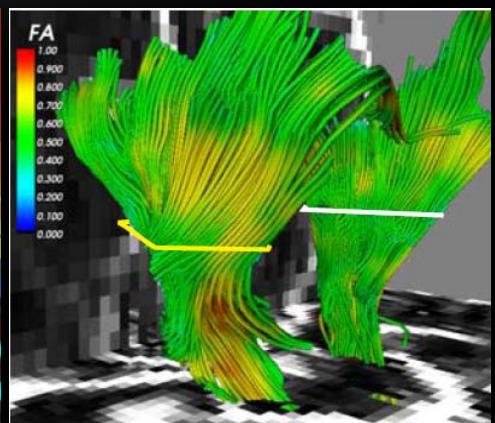
DTI Visualization



Anisotropy Indices



Glyphs



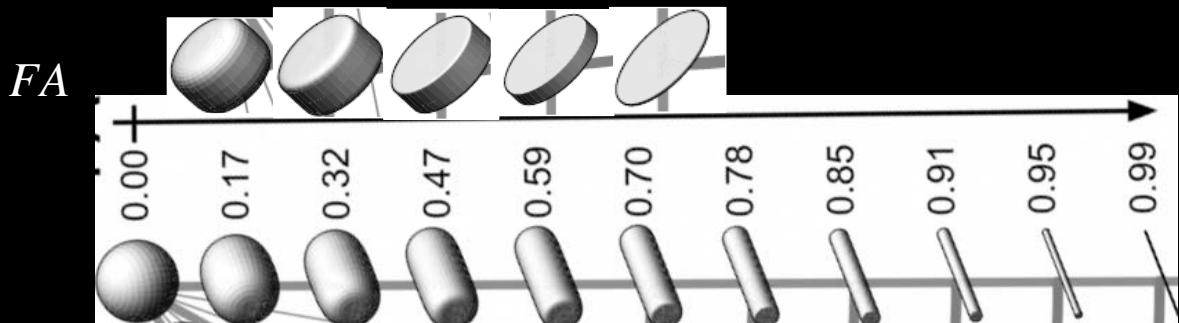
Fiber Tracking

Anisotropy indices

Index that indicates anisotropy

- Fractional Anisotropy

$$FA = \frac{\sqrt{2}}{2} \sqrt{\frac{(\lambda_1 - \lambda_2)^2 + (\lambda_2 - \lambda_3)^2 + (\lambda_1 - \lambda_3)^2}{\lambda_1^2 + \lambda_2^2 + \lambda_3^2}}$$



Geometric Diffusion Measures

Isotropy: $\lambda_1 \approx \lambda_2 \approx \lambda_3$

$$C_s = \frac{3\lambda_3}{\lambda_1 + \lambda_2 + \lambda_3}$$

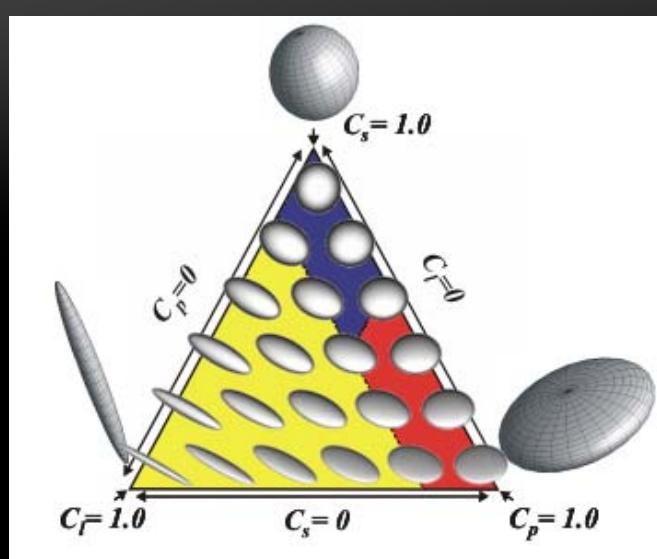
Anisotropy:

Linear $\lambda_1 \gg \lambda_2 \approx \lambda_3$

$$C_l = \frac{\lambda_1 - \lambda_2}{\lambda_1 + \lambda_2 + \lambda_3}$$

Planar $\lambda_1 \approx \lambda_2 \gg \lambda_3$

$$C_p = \frac{2(\lambda_2 - \lambda_3)}{\lambda_1 + \lambda_2 + \lambda_3}$$



$$C_s + C_l + C_p = 1$$

[Westin et al. 97]

Scalar (e.g., Anisotropy index) [Kindlmann et al. 00]

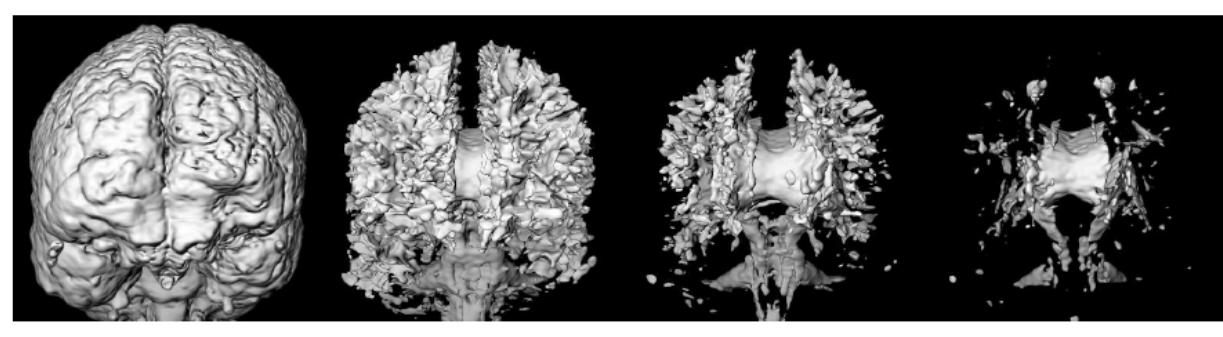


Image from [Vilanova et al. 04]

Anisotropy Indices

There are much more anisotropy indices:

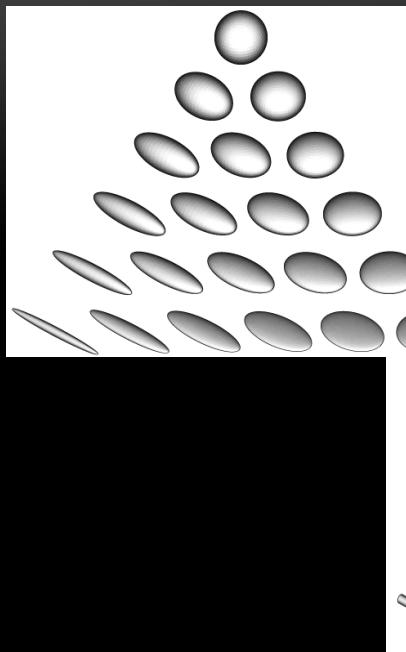
Relative anisotropy (RA), Mean diffusion, etc.

Pros and Cons

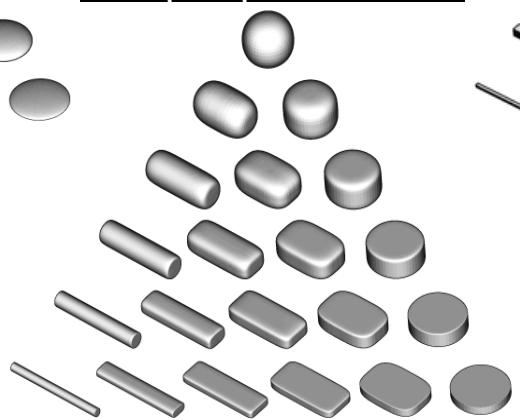
- ✓ “Easy” to visualize
- ✗ Simplification 6D → 1D

That's a hell of a simplification!

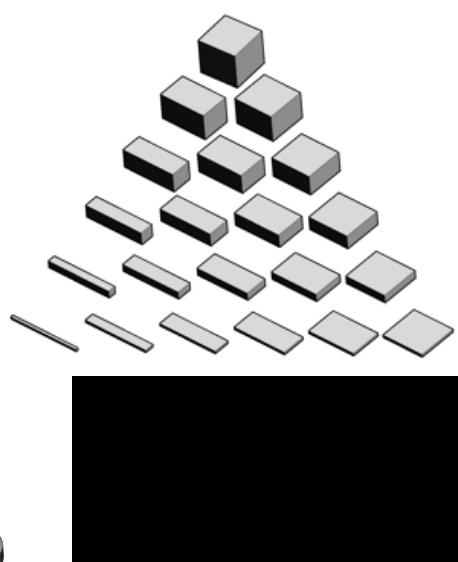
Ellipsoids



Superquadrics



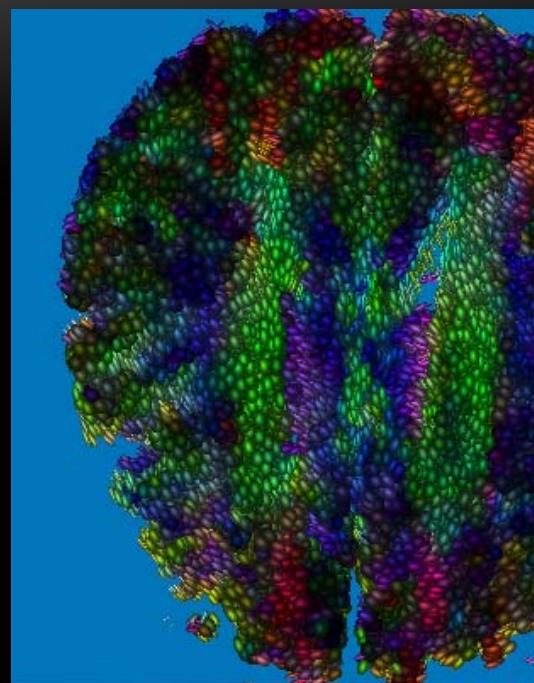
Cuboids



Pros and Cons

- ✓ Shows 6D information
- ✗ Local information
- ✗ Cluttering extended to 3D

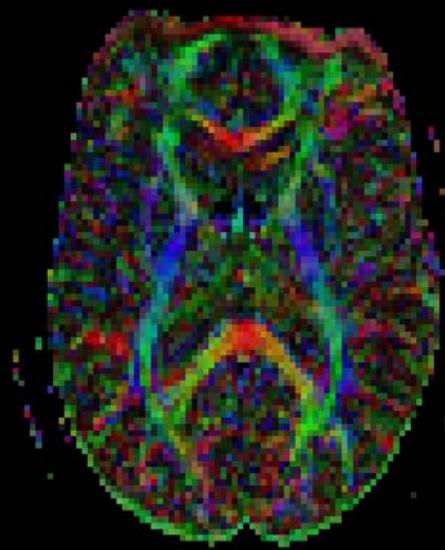
image from [Kondratieva et al. 05]
wwwcg.in.tum.de



$\vec{e}_1 = (x, y, z)$ map to (R, G, B)

Pros and Cons

- ✓ Shows directional information
- ✓ Simple to implement
- ✗ Simplification 6D \rightarrow 3D
- ✗ Difficult to extract fiber information



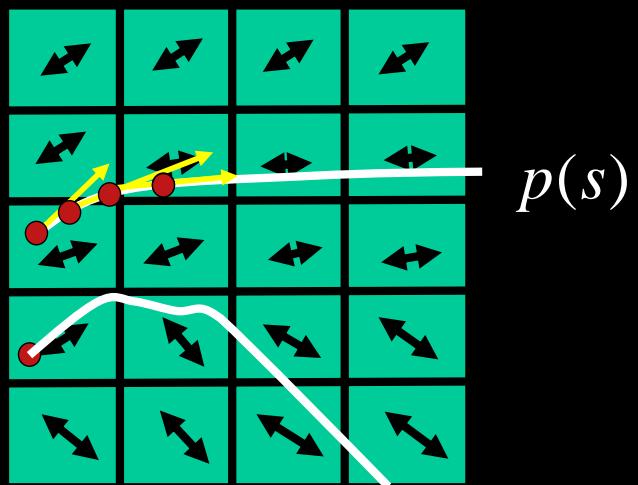
Fiber Tracking *Streamline tracing*

Streamline tracing

$$p(s) = \int \vec{e}_1(p(s))ds \quad p(s) \text{ path with parameter } s$$

Integration scheme

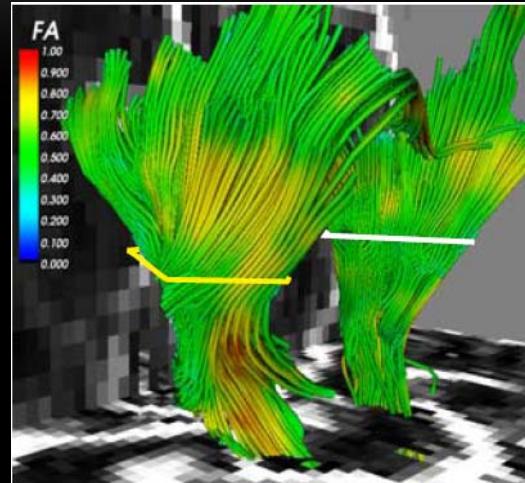
- Euler Forward
- Runge Kutta
- etc.



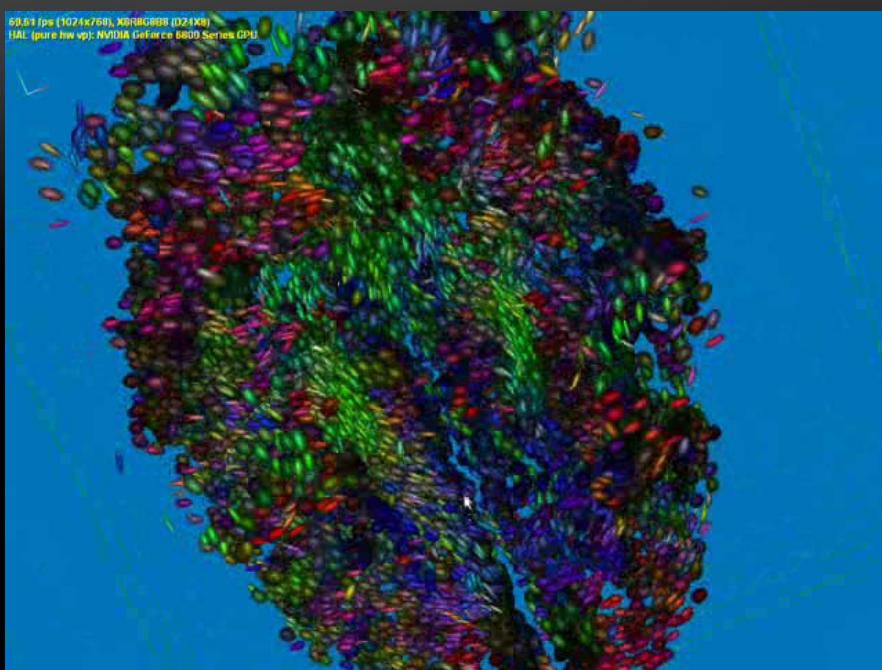
Fiber Tracking Streamline Tracing

Pros and Cons

- ✓ Analogy with fibers
- ✓ Shows global information
- ✗ Simplification 6D → 3D
- ✗ Problems with Crossing
- ✗ Seeding – Region of Interest
- ✗ Cluttering

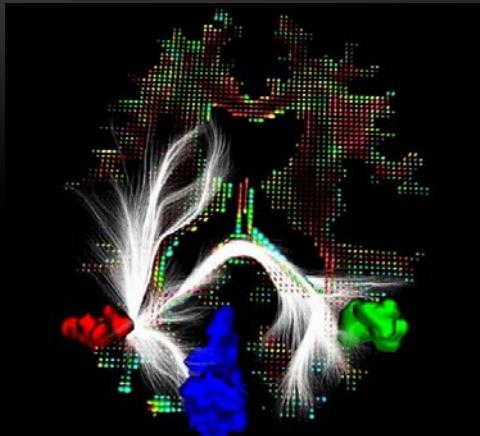


Tracing Glyphs



Video from [Kondratieva et al. 05] wwwcg.in.tum.de

Other Fiber Tracking Techniques

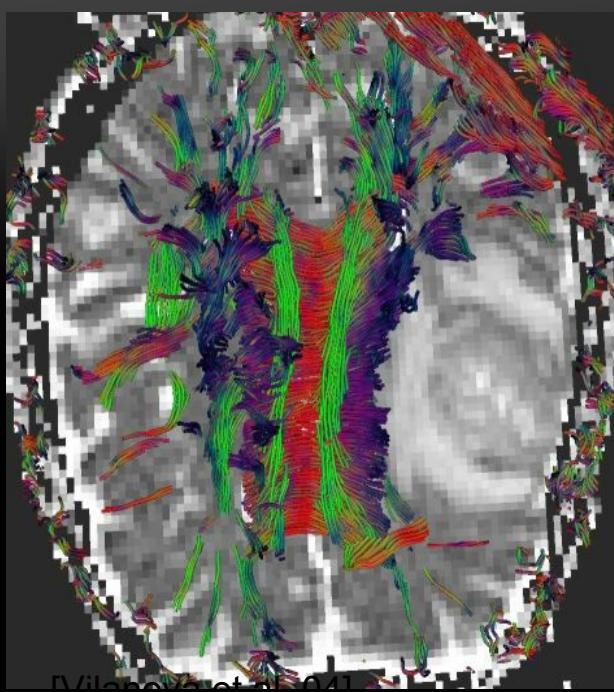


[N.Wotawa et al. 05] [02]

Pros and Cons

- ✓ Analogy with fibers
- ✓ Shows global information
- ✗ Seeding – Initial and end
- ✗ Computational cost
- ✗ Cluttering

Applications



Understanding

- Brain Development
- Brain Injuries
- Ischemic heart
- ...

Diagnosis

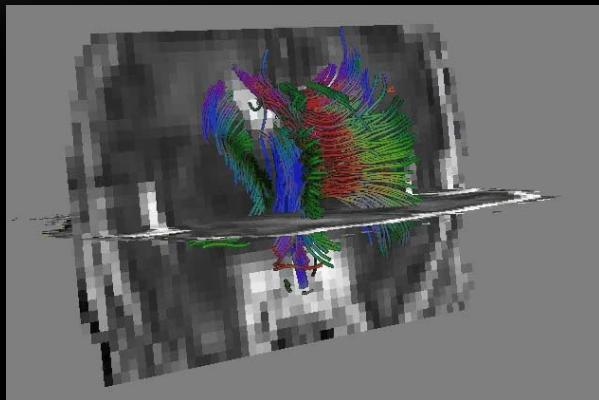
- Epilepsy
- Multiple Sclerosis
- ...

Treatment

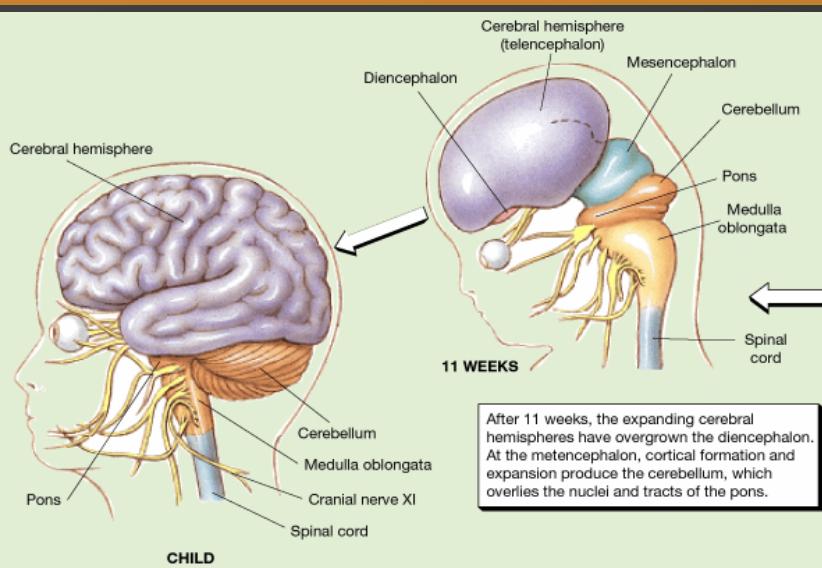
- Tumor resection
- ...

DTI in the newborn brain

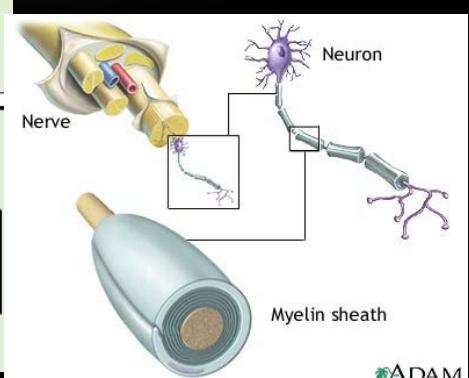
- DTI can reveal detailed anatomy of white matter development.
- Characterization of normal axonal growth of the white matter tracts.
- Understanding the extensive inhomogeneity of white matter injuries (e.g., hypoxic-ischemic regions)
- Reference standards for diagnostic radiology of premature newborns.
- Early detection can improve treatment



Human brain development



Picture from Prentice Hall -
cwx.prenticehall.com



Brain myelination starts with 30 weeks of conception and it is not completed until the age 20-30

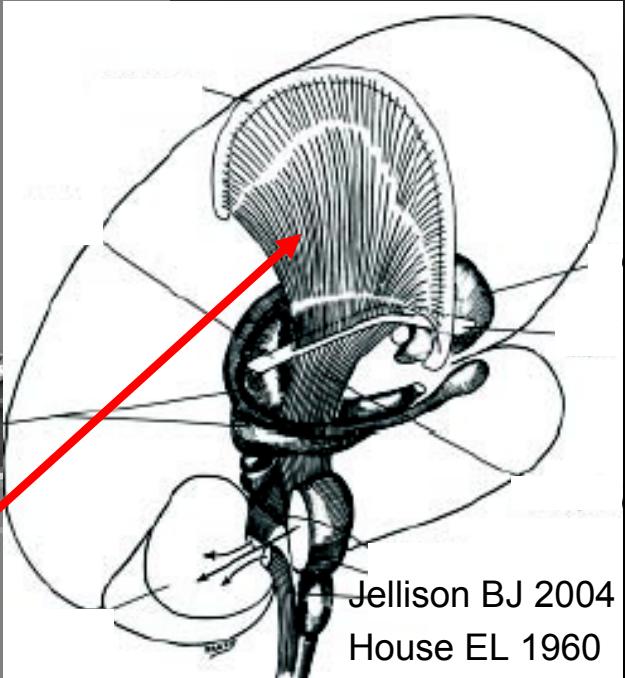
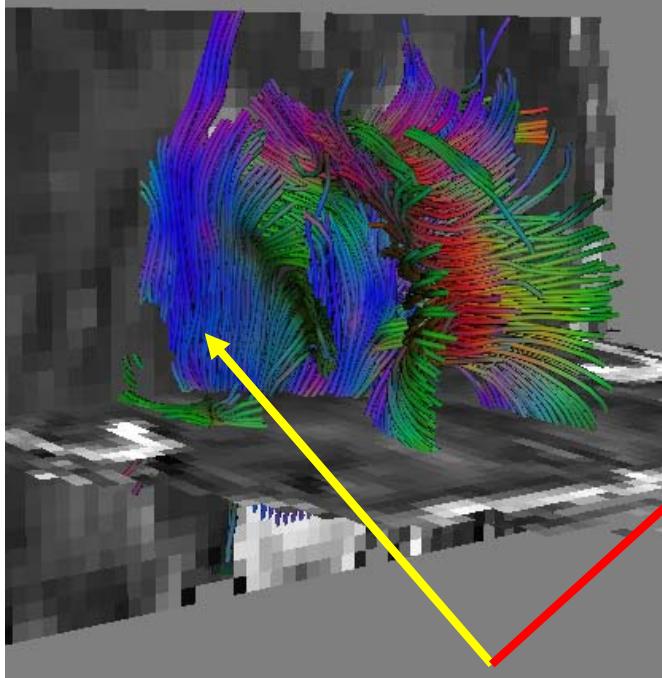
Acquisition Difference with Adults:



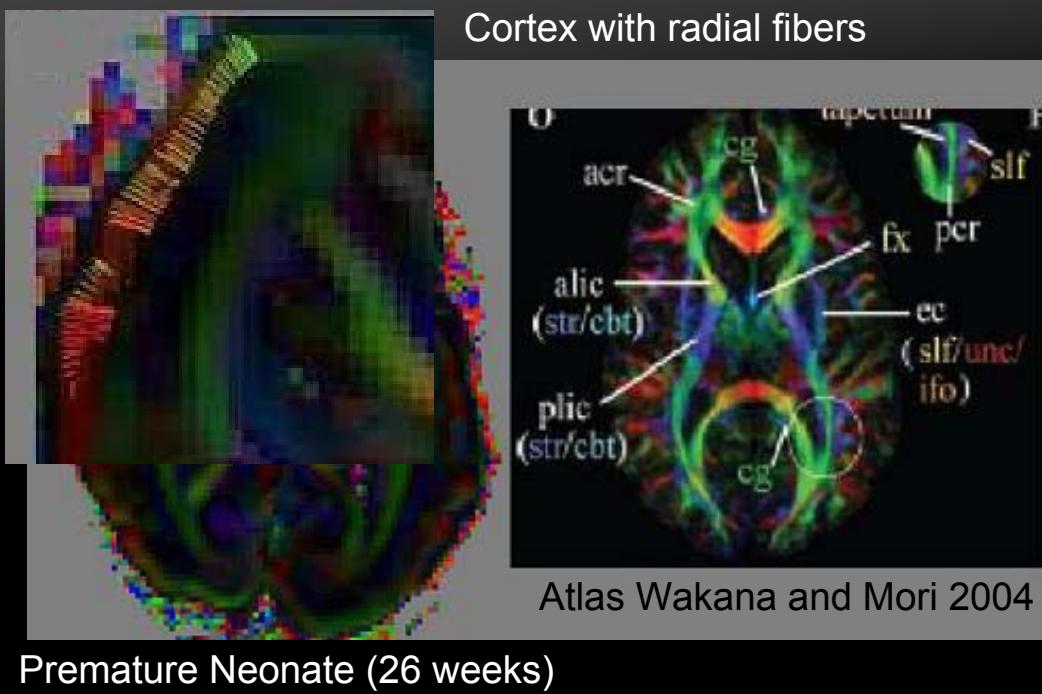
- Fibers are less myelinated → less anisotropy → lower signal intensity
- Motion artifacts can play a larger role (scan within 4 minutes full-term newborns)
- The size of the pre-term (and neonatal) brain is smaller than of an adult.
Voxel contains more structures than in an adult.
- The signal strength decreases if the voxel size decreases.

Visualization DTI: fiber tracking

Full term newborn at day 6



Example: Adult vs.Premature Newborn

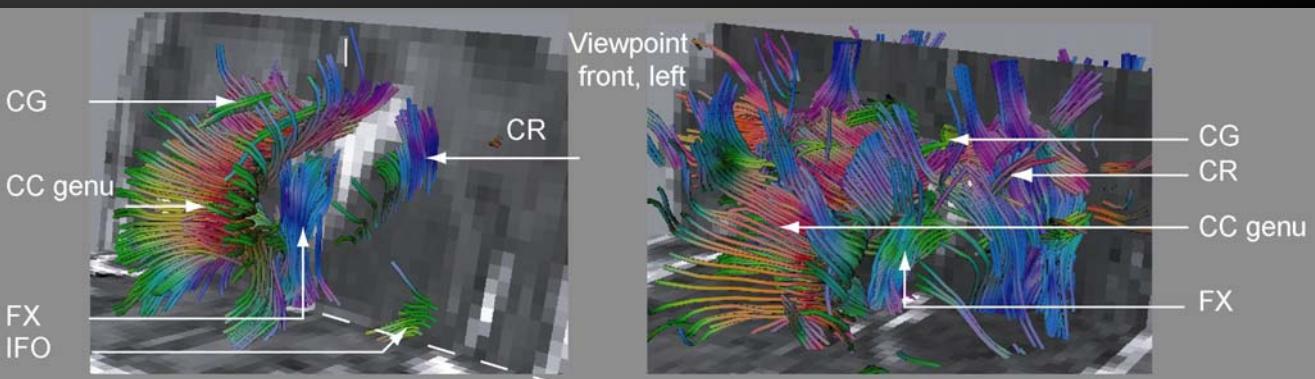


Premature Neonate (26 weeks)

Results: normal newborns (follow ups)

birth

3 months

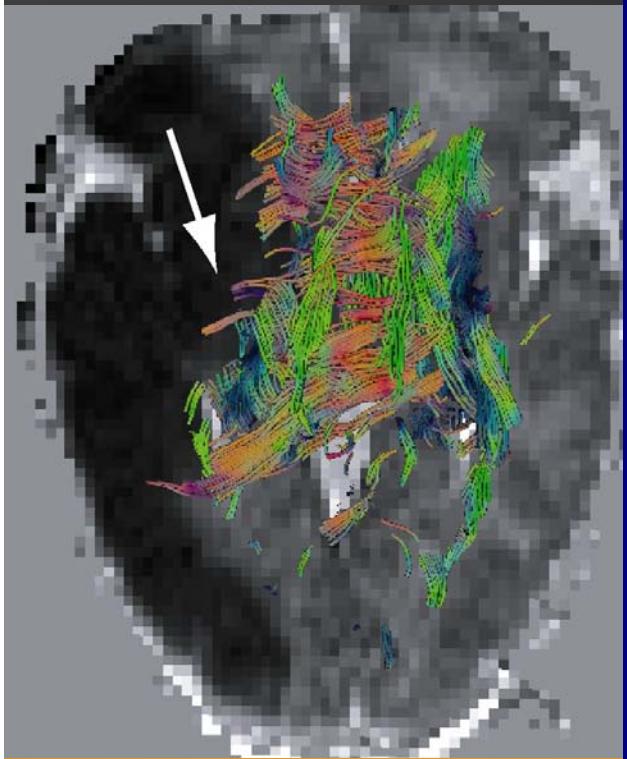


Which structures are developing and how?
Quantification?

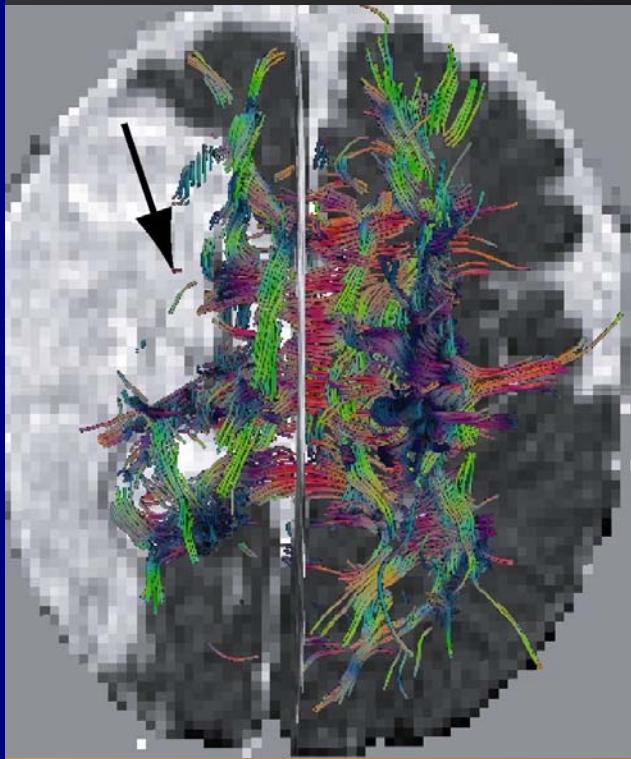
Results: newborns with pathology

Vis08
VIS • INFOVIS • VAST

birth

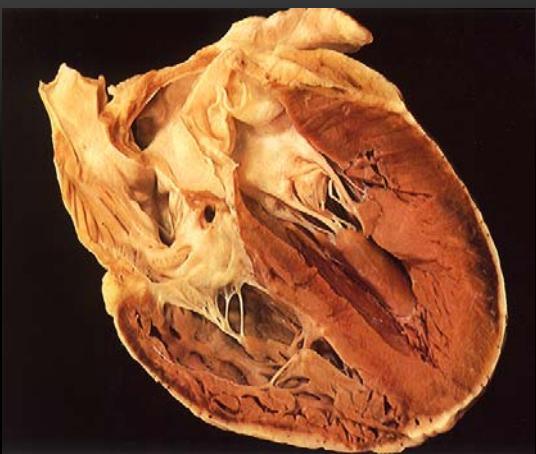


3 months

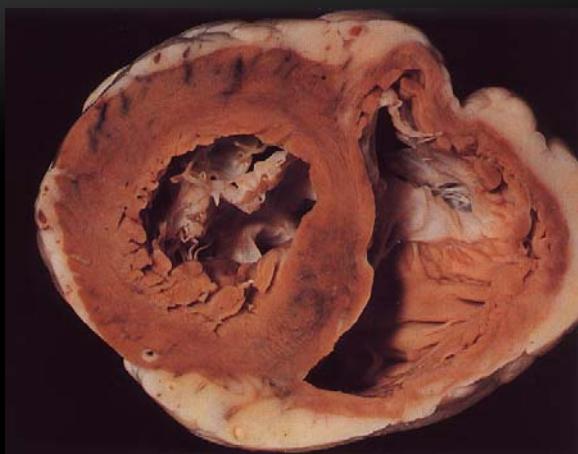


Heart DTI visualization

Vis08
VIS • INFOVIS • VAST

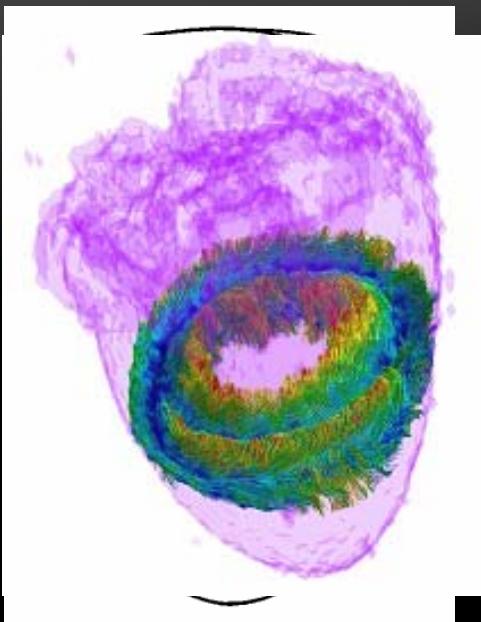


Sagittal section through the heart

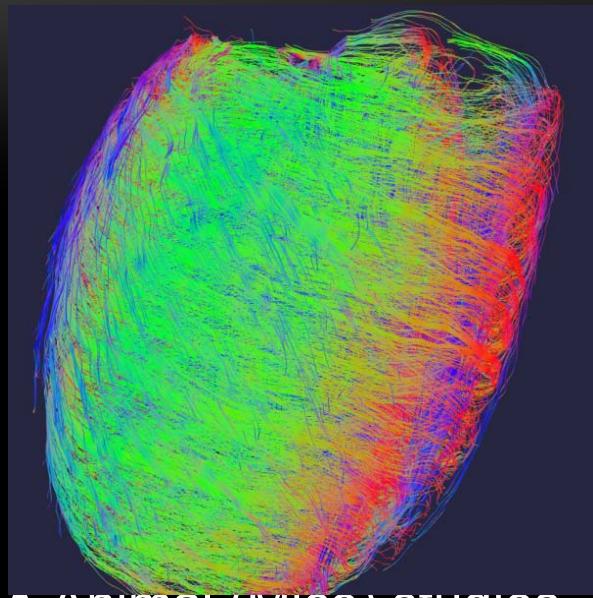


Short-axis section through the heart

[Anderson, 1980]



Model of Left Ventricle, depicting helix angle [Bovendeerd, 1992]

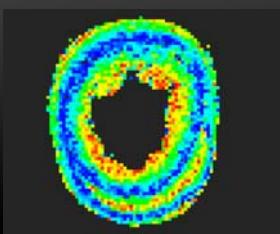


- Animal (mice) studies

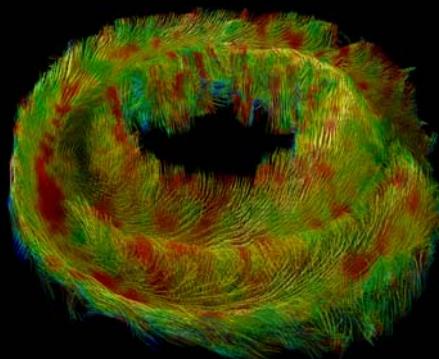
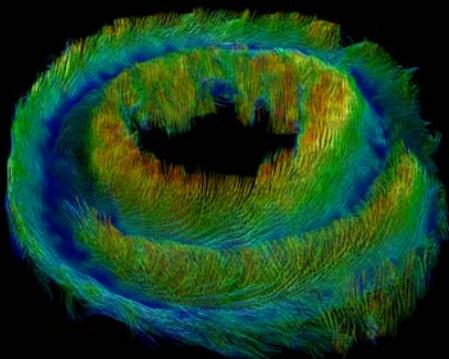
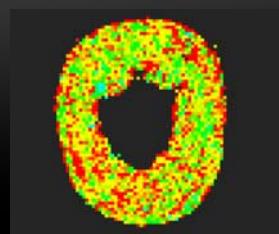
Visualization of the heart

Hue color mapping

Helix Angle



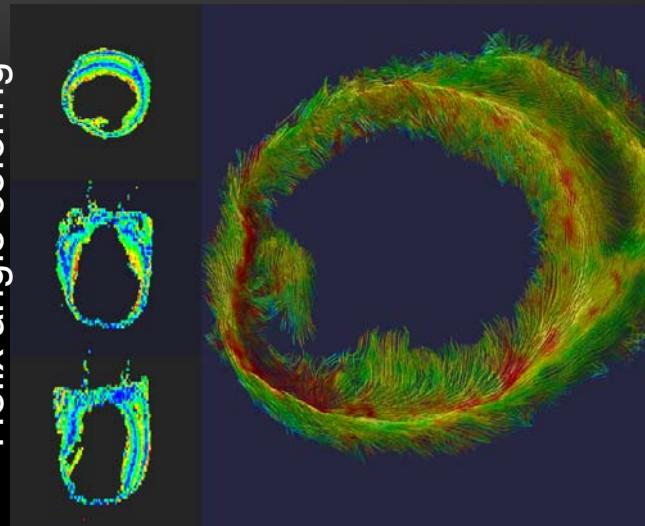
Hue color mapping
Fractional Anisotropy



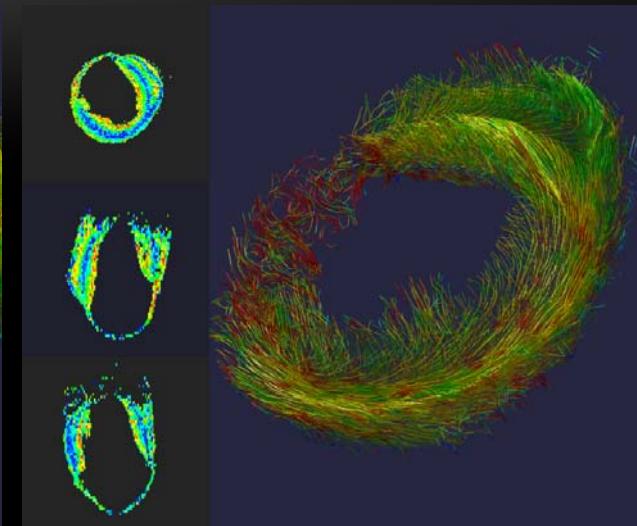
Illuminated lines + Shadows

Fibers in a slice of ischemic mouse hearts

Helix angle coloring



7 days
after infarct



28 days
after infarct

Ischemic hearts

In ischemic areas:

- Heart-wall becomes thinner
- FA becomes higher (this was unexpected)
- More random fiber orientations

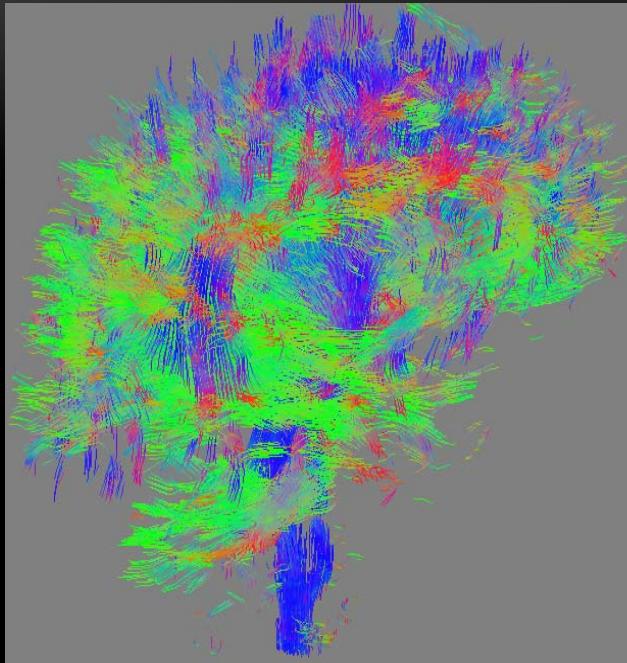
Conclusion: High FA and random fiber orientation
probably caused by collagen fibers

Seed point definition

- Region of interest
 - Biased
 - Not reproducible
 - Miss information
- Whole volume
 - Cluttering

Individual “fibers” are of no interest

Bundles structures are of interest



Fiber Bundle

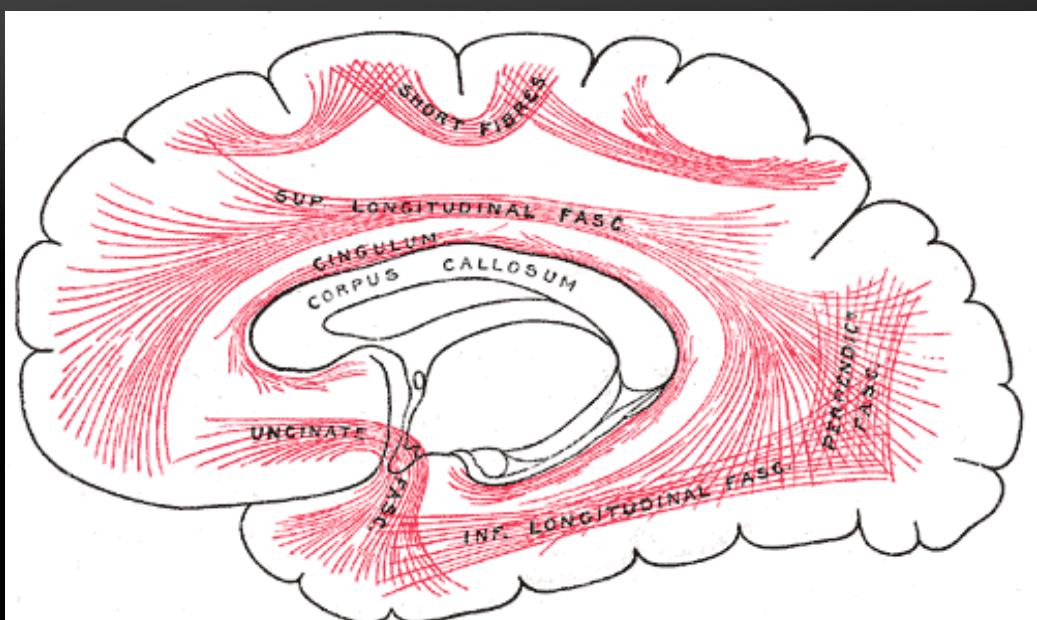


Image from Brun et al. 2003

Fiber Clustering

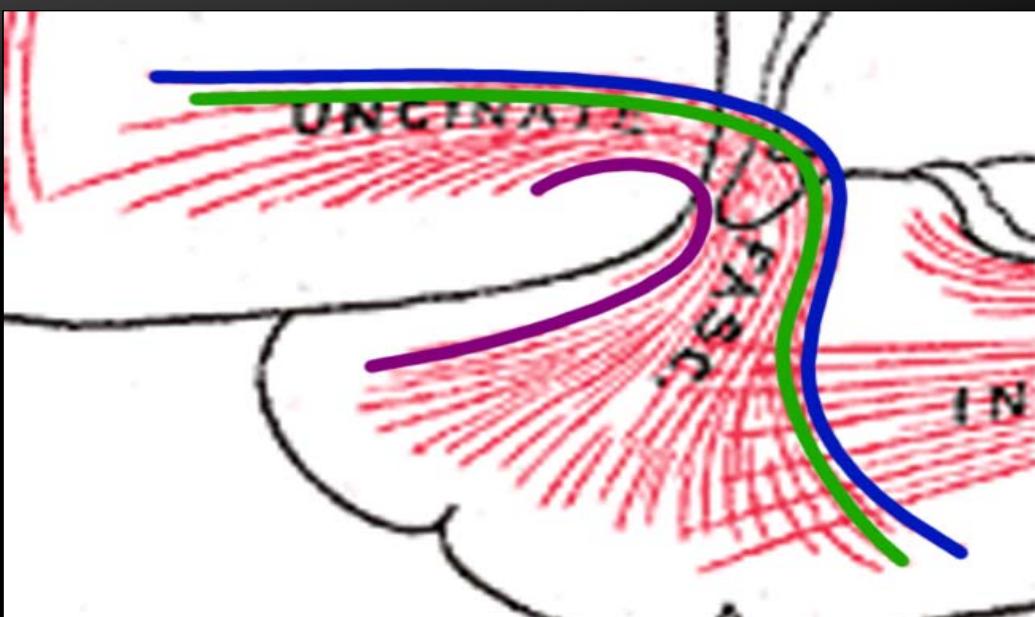
Group fibers together that are *similar*

Form fiber bundles that are meaningful

Two problems:

- How to measure similarity between fibers?
- How to define the groups of fibers?

Fiber Bundle Properties



A lot of possible combinations

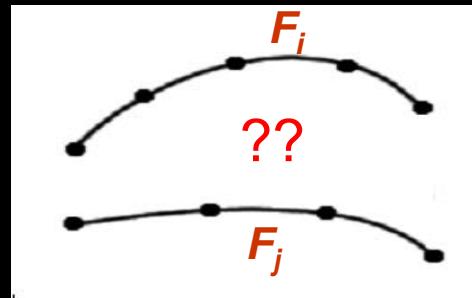
Ding et al. 01, Shimony et al. 02, Zhang et al. 02, Brun et al. 03, Brun et al. 04, Corouge et al. 04, etc.

There are a lot of similarity measures :

- Mean of closest points distance (Corouge et al. 04)
- Closest point distance (Corouge et al. 04)
- Hausdorff distance (Corouge et al. 04)
- End points distance (Brun et al. 03)
- ...

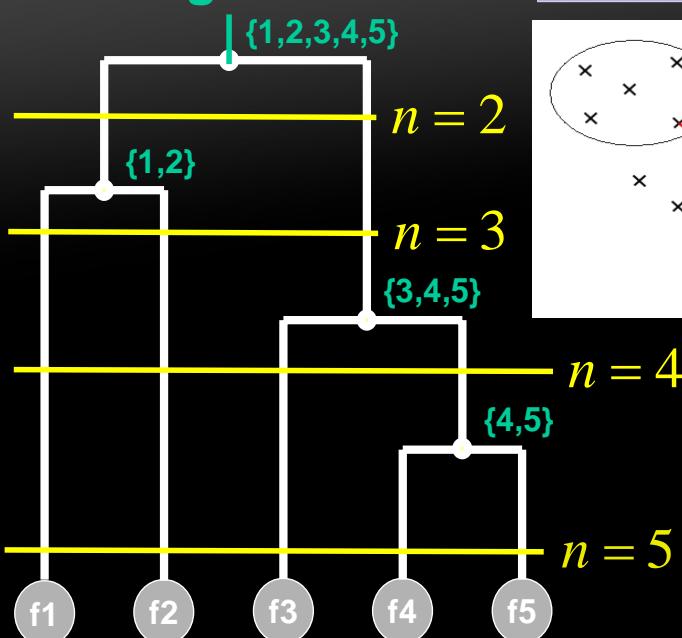
There are a lot of clustering algorithms:

- Hierarchical (Zhang et al. 02)
- Fuzzy c-means (Shimony et al. 02)
- Spectral clustering (O'Donnell and Westin 05)
- Shared nearest neighbor (Moborts et al. 05)
- ...

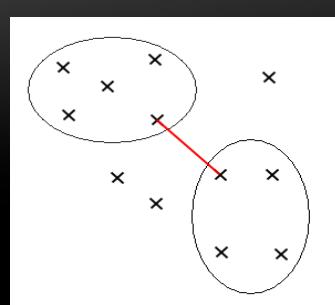


Example: Hierarchical Clustering

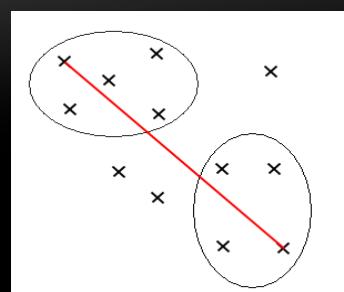
Dendrogram



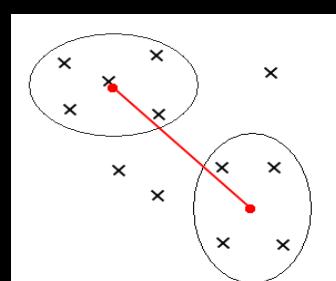
Single Link (HSL)



Complete Link (HCL)



Weighted Average(HWA)

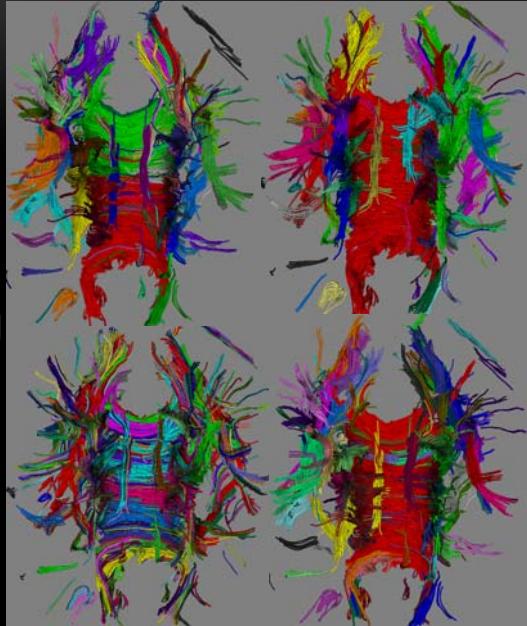


How do we know that ...

- ... a method is better than the other?
- ... a similarity measure is better than another?
- ... there is not a parameter setting giving better results?

Validation [Moberts et al. 05]

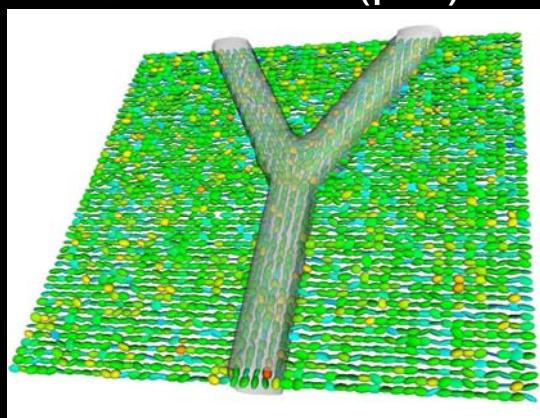
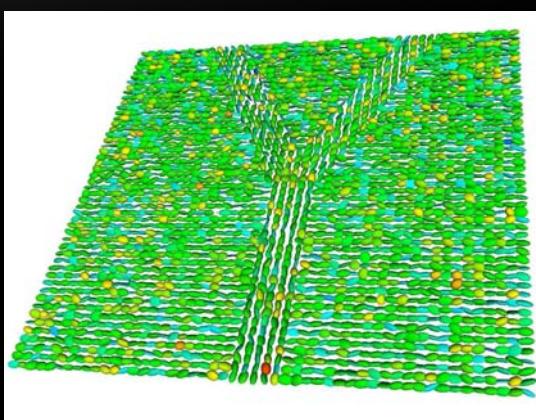
- Ground truth
- Comparison framework



Diffusion Tensor Imaging Segmentation

Fiber clustering depends on the fiber tracking algorithm and its parameter settings.

Can we directly segment the tensor fields (pdf)?

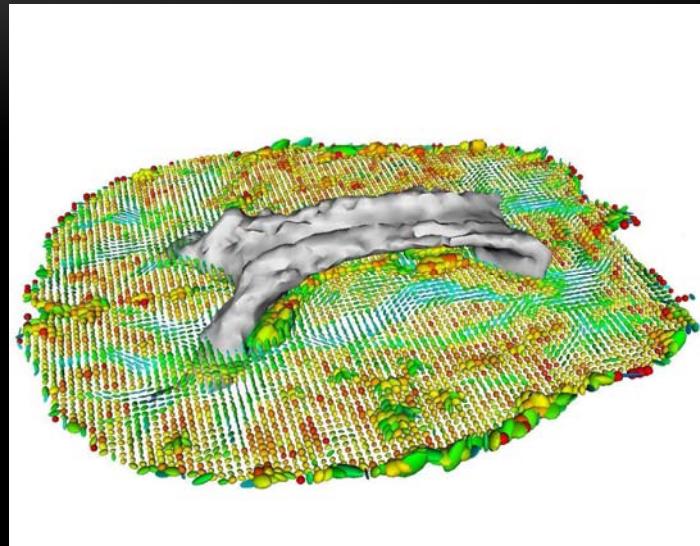


Images from [Lenglet et al. 04]

Diffusion Tensor Imaging Segmentation

There exist several segmentation techniques for scalars:

- **Thresholding:** ordering of the tensors
- **Region based:** definition of homogeniety (e.g. Ziyang et al 06, Bartesaghi and Nadar 06)
- **Edge based:** definition of gradient in the tensor field
- **Defomable Models:** definition of forces and energies based on the tensor field (e.g., Lenglet et al 06, Wang et al. 05, Schultz et al. 06)
- ...



Corpus callosum segmentation
using level sets technic
Images from [Lenglet et al. 04]

Tensor similarity or distance

You want to group diffusion tensors that are similar.

Given tensor A and B, How similar (different) are they?

- Linear Algebra- tensor is a 6D vector . Example:

$$d_{L2}(A, B) = \sqrt{\sum_{i=1}^3 \sum_{j=1}^3 (A_{ij} - B_{ij})^2}$$

- Riemannian geometry- geodesic distance in the space of positive definite matrices.

$$d_g(A, B) = N(A^{-\frac{1}{2}} B A^{-\frac{1}{2}})$$

$$N(D) = \sqrt{\sum_{i=1}^3 (\log(\lambda_i^D))^2}$$

Approximation Log-Euclidean distance

$$d_{LE}(A, B) = \sqrt{\text{tr}((\log(A) - \log(B))^2)}$$

- Probability density functions (pdf) – overlap of the pdf using A and B as covariant matrices of Gaussians

- Kullback-Leibler (KL) distance

$$d_{KL}(A, B) = \frac{1}{2} \sqrt{\text{tr}(A^{-1}B + B^{-1}A) - 2n} \quad n \text{ is } 3$$

- Class separability Bhattacharyya bound

$$S_{Bhat}(A, B) = e^{-\frac{1}{2} \ln \left(\frac{\det(AB)/2}{\sqrt{\det(A)\det(B)}} \right)}$$

- Anisotropy Indices – use anisotropy indices FA, CI or any combination of those
- Angular differences – Use the angular difference between the main eigenvectors (dot product)

DTI Segmentation

- What measure or method to use depends on the problem (e.g., bundle)
- Segmentation is an active field of research for scalar fields. Extension to tensor fields is a challenge
- These methods use the full information of the tensor and can be more robust and reproducible than fiber clustering techniques
- No much has been done in this field yet

- Diffusion Tensor Imaging data
- DTI Visualization techniques
- Applications: newborn and ischemic heart
- Fiber clustering
- Diffusion tensor field segmentation

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