### Diffusion Modeling in BrainSuite

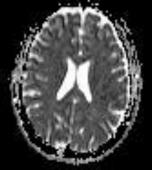
Justin P. Haldar



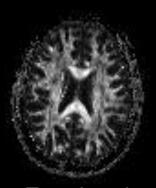
### Motivation

Diffusion MRI provides unique insight into tissue microstructure

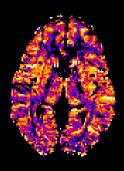
Quantitative measures



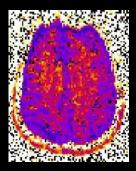
Apparent
Diffusion
Coefficient



Fractional Anisotropy

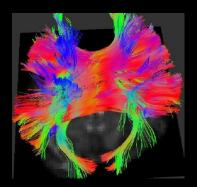


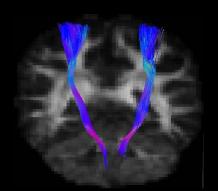
Anomalous Exponent

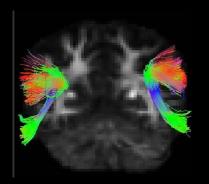


**Kurtosis** 

Connectivity







# Diffusion Imaging (in general)

- 1. Acquisition
- 6+ Directions
- HARDI
- Multi-Shell
- DSI
- ...

- 2. Preprocessing
- Registration
- Distortion Correction
- Outlier Removal

- 3. Modeling
- DTI
- Multi-DTI
- ODF
  - FRT
  - FRACT
  - Nonlinear Methods
- NODDI, DKI, ...

- 4. Tracking
- Deterministic
- Probabilistic

- 5. Analysis
- Visualization
- Connectivity Analysis
- ROI Analysis

# Diffusion Imaging (in BrainSuite)

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- 6+ Directions
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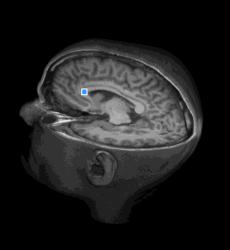
- 3. Modeling
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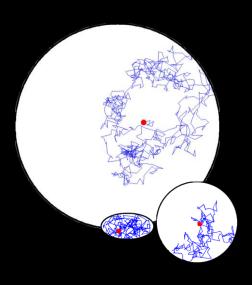
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- Probabilistic

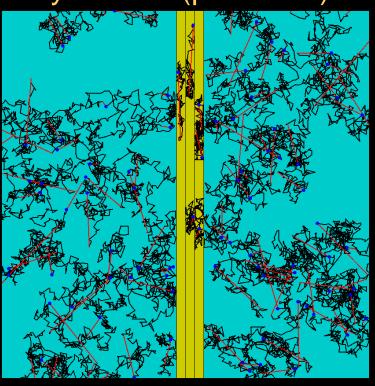
- 5. Analysis
- Visualization
- Connectivity Analysis
- ROI Analysis

## Data Sampling

- Diffusion data is sampled from a 6D space
  - 3D Spatial
    - Where in the brain am I located? (mm scale)
  - 3D Diffusion
    - Where do water molecules typically move? (µm scale)

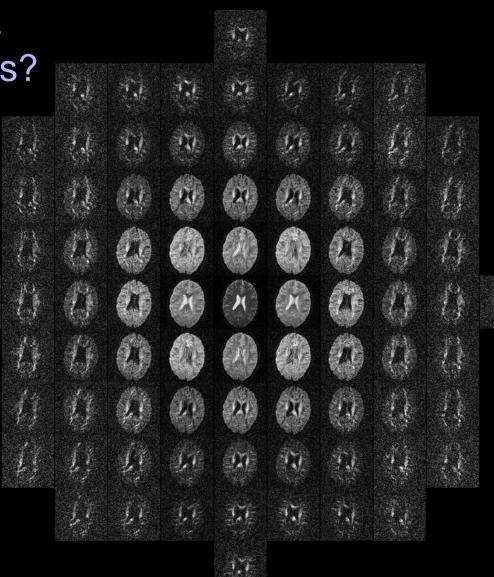






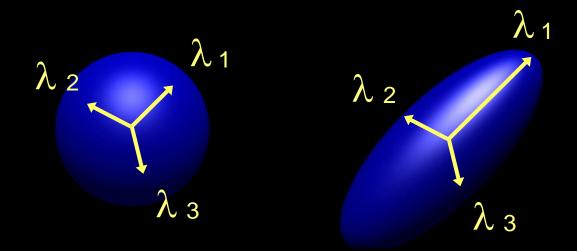
### Diffusion Data is Complicated

- Sampling in 6D = time
- How to interpret results?
- Solution:
  - Models!
  - In BrainSuite:
    - Diffusion Tensor Imaging (DTI)
    - Orientation Distribution Functions (ODFs)
      - Funk Radon Transform (FRT)
      - Funk Radon and Cosine Transform (FRACT)

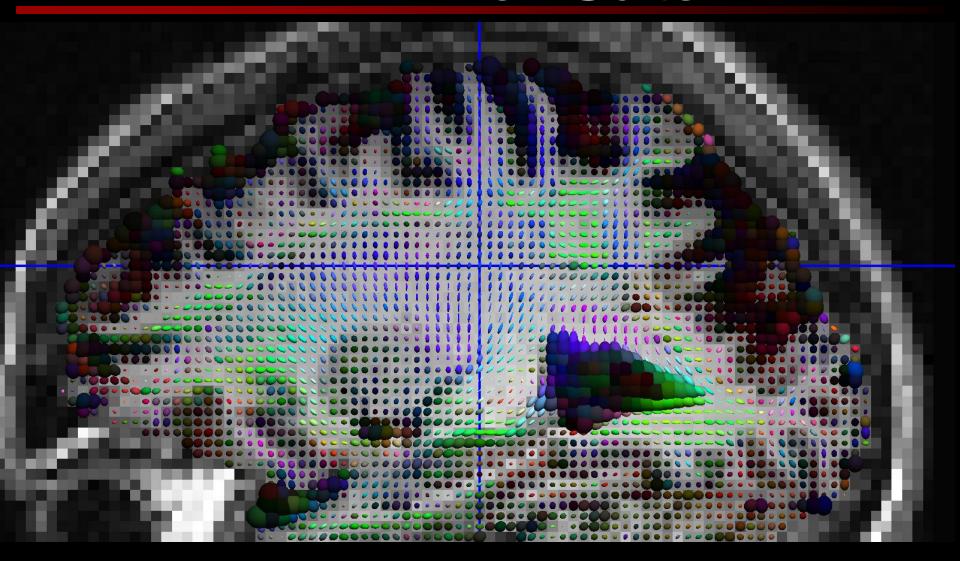


#### DTI

- Diffusion Tensor Imaging:
  - A simple model with 7 degrees of freedom
    - 3 orthogonal directions
    - 3 diffusivities along each direction
    - 1 baseline image intensity



# DTI in BrainSuite

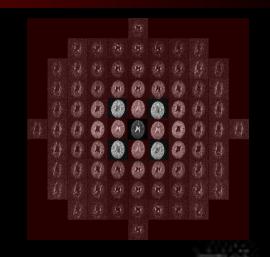


### Benefits of DTI

- Only need to sample 7 images
  - In practice ~30 is preferred
  - Sampling is very flexible
- Quantitative information:
  - Mean Diffusivity
    - $(\lambda_1 + \lambda_2 + \lambda_3)/3$
  - Fractional Anisotropy

$$\sqrt{\frac{3}{2}} \frac{\sqrt{\left(\lambda_1 - \bar{\lambda}\right)^2 + \left(\lambda_2 - \bar{\lambda}\right)^2 + \left(\lambda_3 - \bar{\lambda}\right)^2}}{\sqrt{\lambda_1^2 + \lambda_2^2 + \lambda_3^2}}$$

- Tissue Orientation
  - Enables tracking



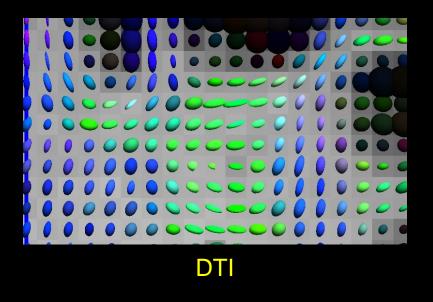


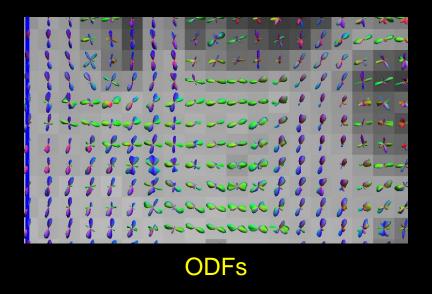


### Limitations of DTI

- Model is almost never valid
  - (But it still provides good biomarkers for microstructure)

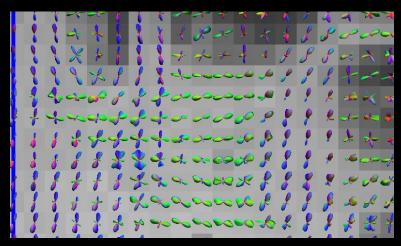
Cannot model multiple orientations in a single voxel



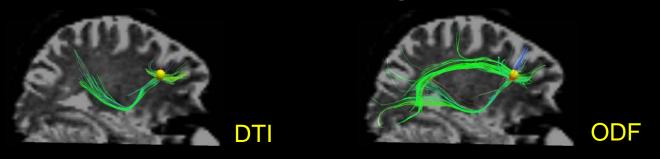


### Orientation Distribution Functions

 Give the ability to model multiple orientations in a single voxel

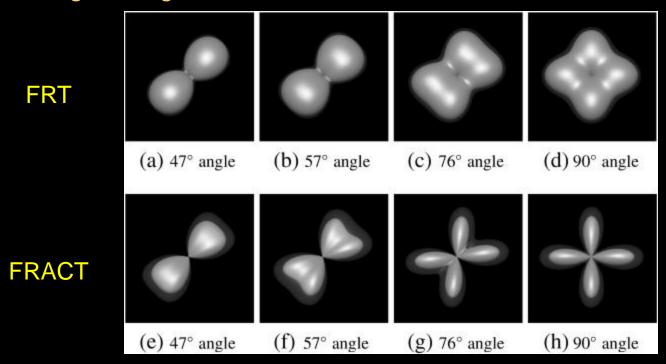


 Enables better tracking in complicated regions with multiple fiber crossings



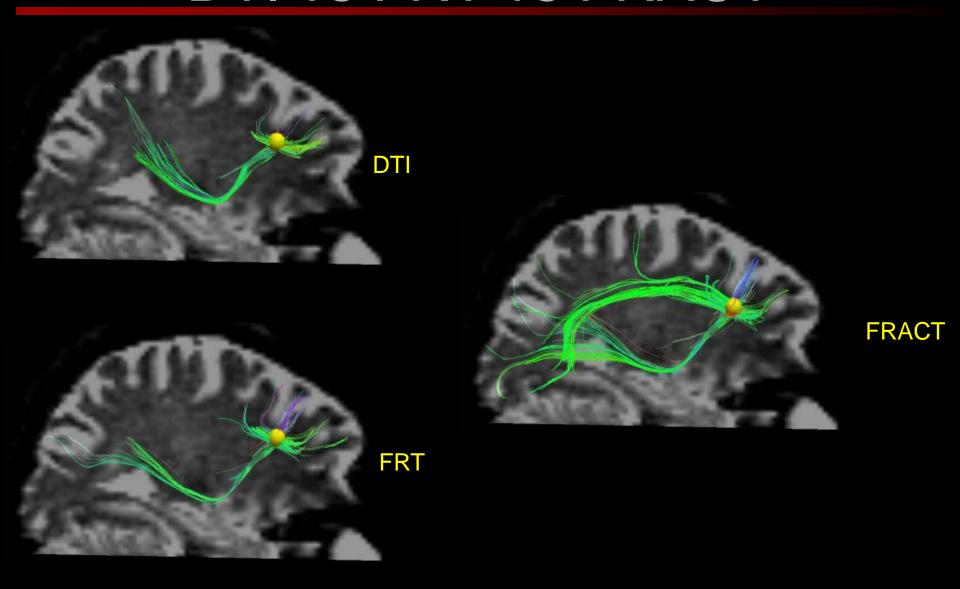
### FRT vs. FRACT

- BrainSuite Provides two ODF methods:
  - Funk Radon Transform [1]
    - Most prevalent ODF method in other software
  - Funk Radon and Cosine Transform [2]
    - Higher angular resolution



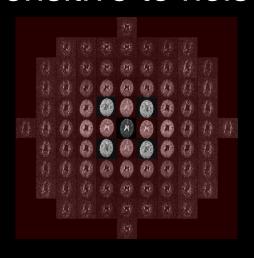
[1] Tuch, Magn Reson Med 52:1358-1372, 2004.[2] Haldar and Leahy, Neurolmage 71:233-247, 2013.

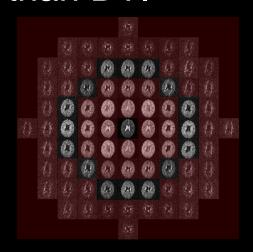
# DTI vs FRT vs FRACT



### Limitations of FRT+FRACT

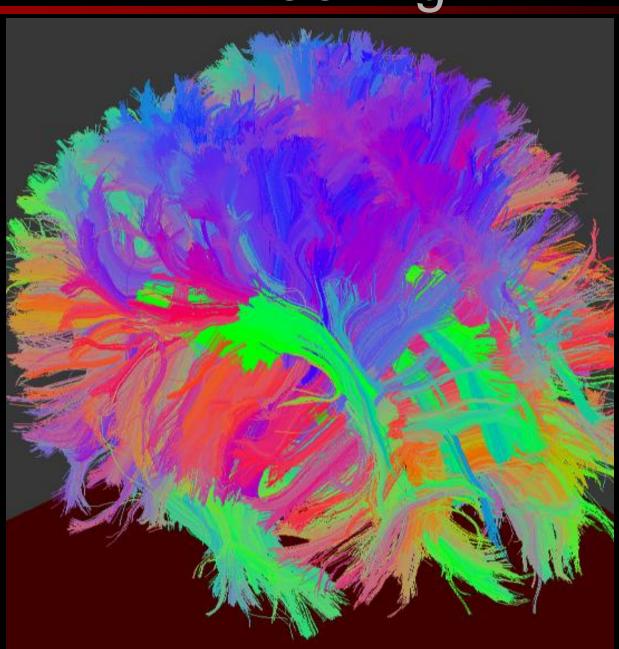
- Requires more data than DTI
  - 64+ directions recommended (often more is needed)
- Works only for data on the sphere
  - better with high b-value data
- More sensitive to noise than DTI





DTI

# Tracking

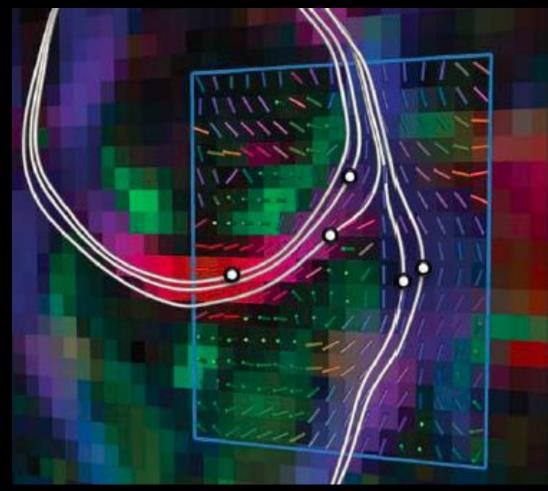


# Deterministic Tractography

Start at seed locations

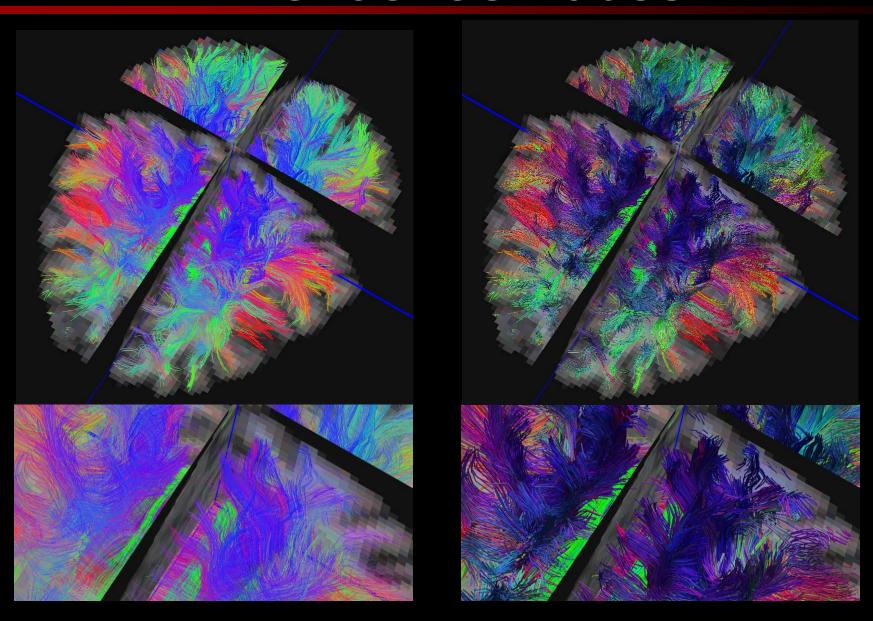
 Follow major orientation

Know when to stop

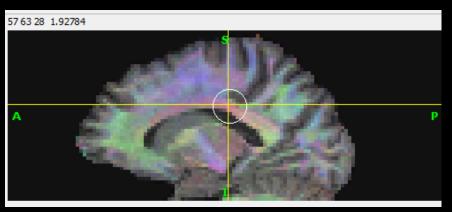


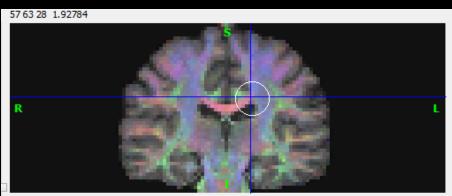
Tournier 2011

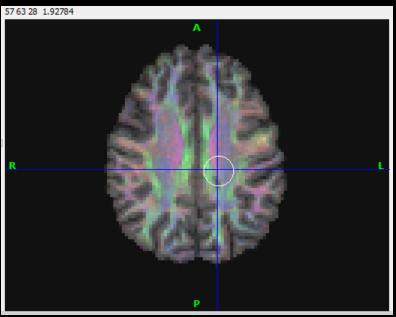
# Render as Tubes

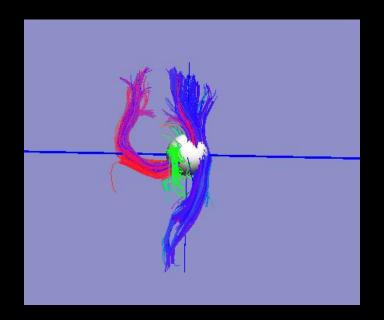


# Sphere Filtering

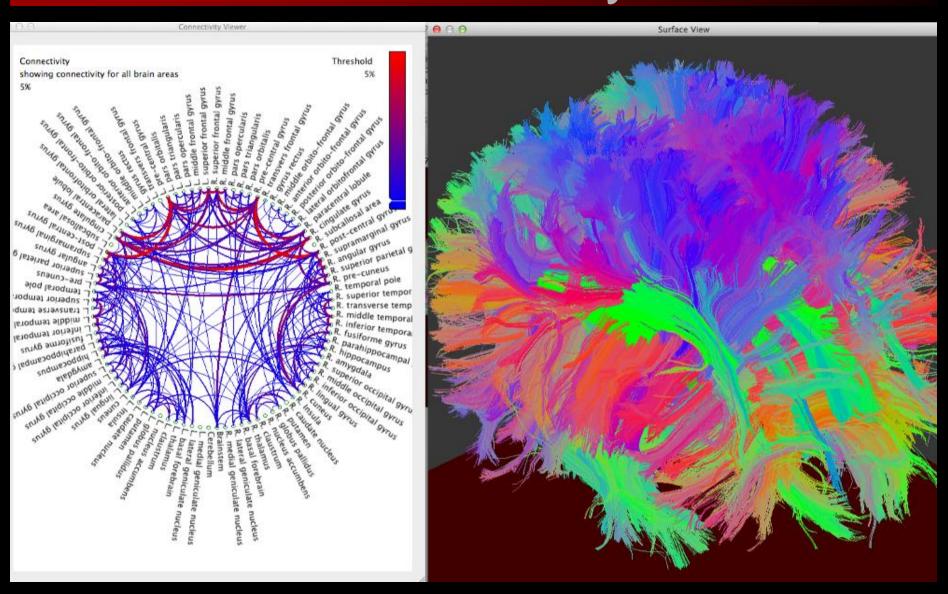






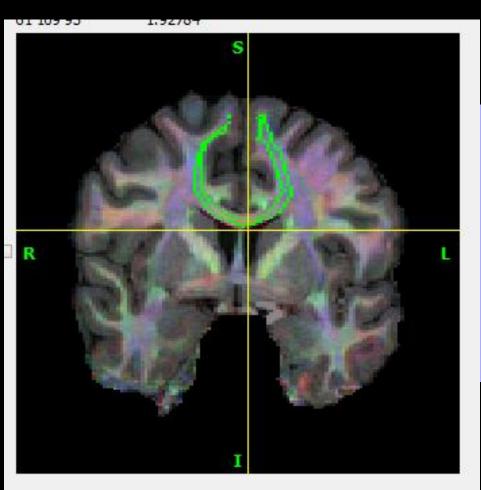


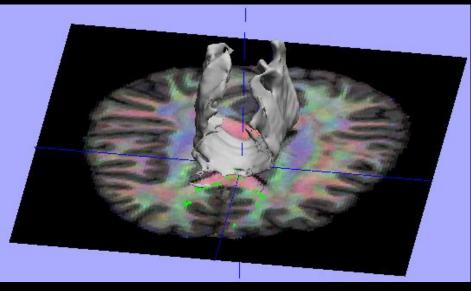
# Connectivity



### Track-Based ROI Tool

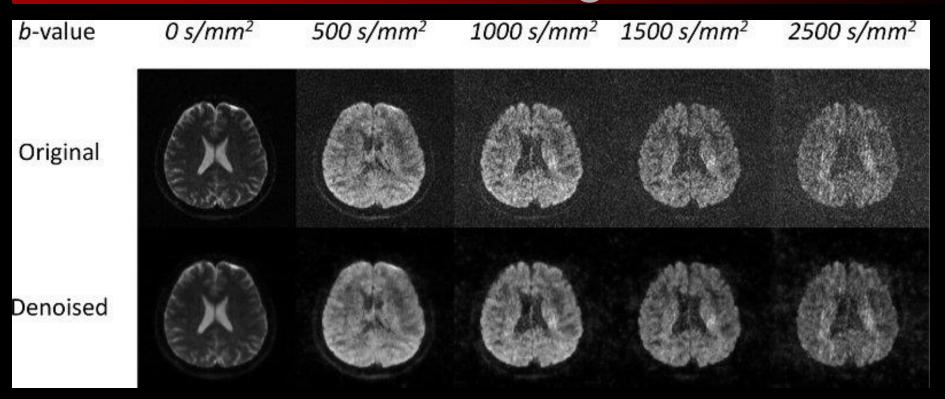
Select voxels based on tracks





http://brainsuite.org/processing/additional-tools/

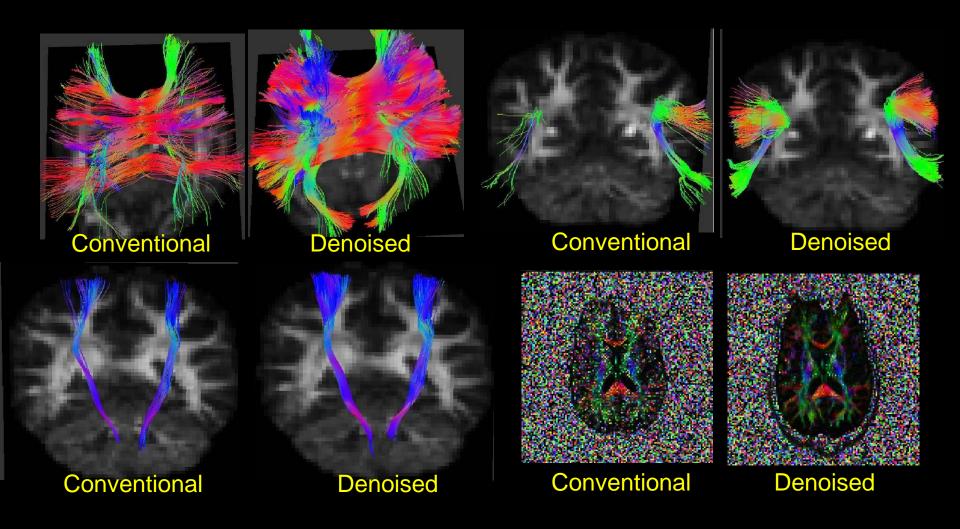
### Denoising



- [1] Haldar, Wedeen, Nezamzadeh, Dai, Weiner, Schuff, Liang, *Magn Reson Med* 69:277-289, 2013.
- [2] Varadarajan and Haldar, IEEE ISBI, 2013, pp. 708-711.

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### Biases and Errors

- Diffusion is powerful!
- But make sure you know how to interpret the results!

# Twenty-five Pitfalls in the Analysis of Diffusion MRI Data<sup>†</sup>

NMR Biomed. 2010; 23: 803-820

Derek K. Jones<sup>a</sup>\* and Mara Cercignani<sup>b</sup>

Challenges and limitations of quantifying brain connectivity in vivo

with diffusion MRI

#### **Derek K Jones**

Imaging Med. (2010) 2(3), 341-355

White matter integrity, fiber count, and other fallacies: The do's and don'ts of diffusion MRI

NeuroImage 73 (2013) 239-254

Derek K. Jones a,b,\*, Thomas R. Knöschec, Robert Turnerc