

**Scuola Superiore di Neurologia **

**V CORSO**

***Neuroimmagini nella Malattia di Parkinson e Parkinsonismi***

**Genova, 21-22 febbraio 2017**

**Accademia Nazionale di Medicina - Via M. Piaggio 17/6 - Genova**

## **DIFFUSION MRI**

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Milan, Italy

# DIFFUSION MRI

## Outline of the presentation

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- ✓ Basic principles
- ✓ Parkinson's disease
- ✓ Progressive supranuclear palsy
- ✓ Corticobasal syndrome
- ✓ Multiple system atrophy
- ✓ Differential diagnosis

# DIFFUSION MRI

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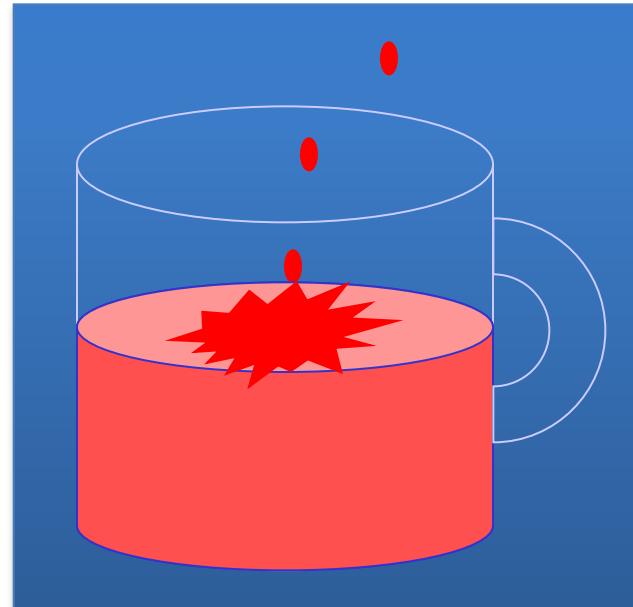
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# DIFFUSION MRI

## Basic principles

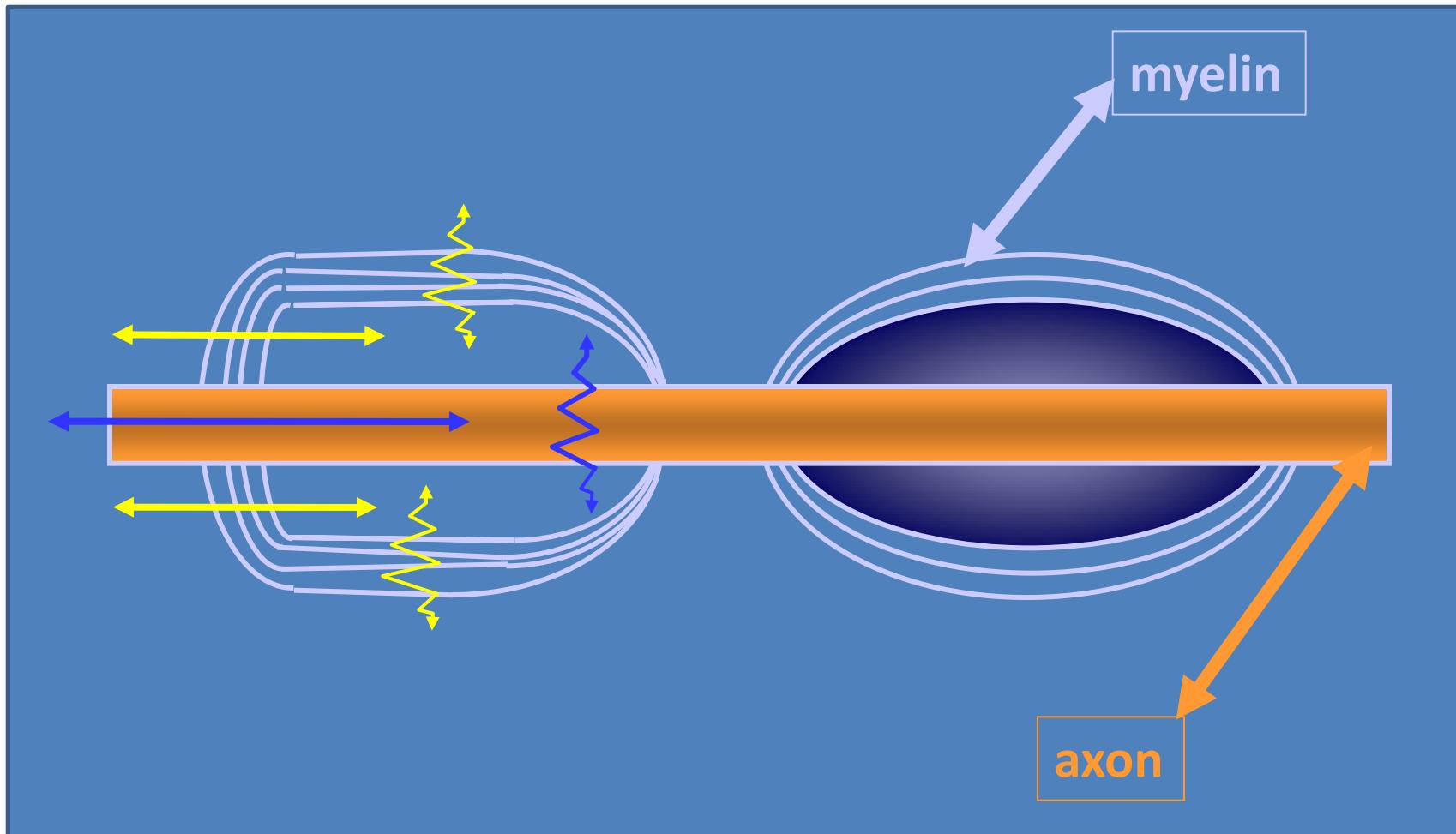
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Diffusion results from a microscopic random motion known as Brownian motion

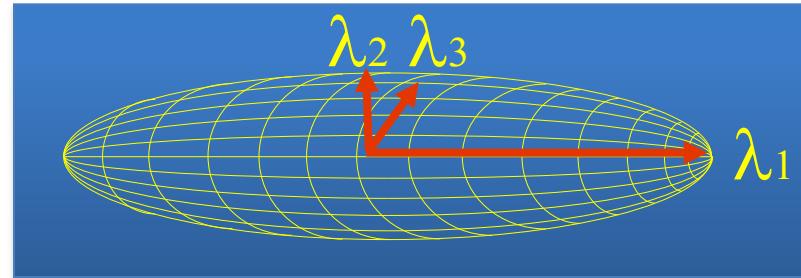
# DIFFUSION MRI

## Diffusion anisotropy in white matter

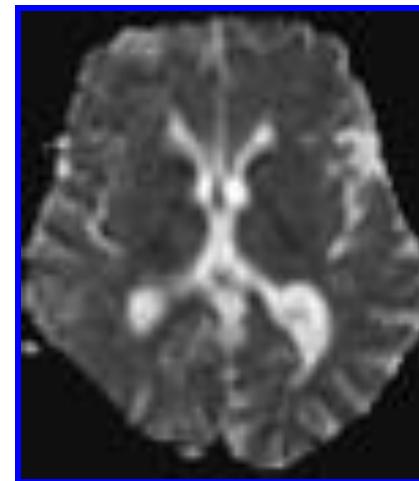
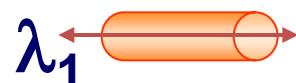
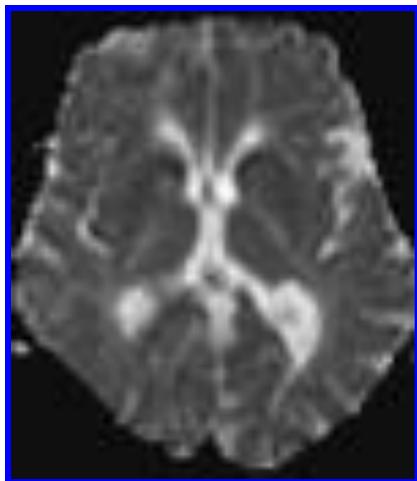


# DIFFUSION MRI

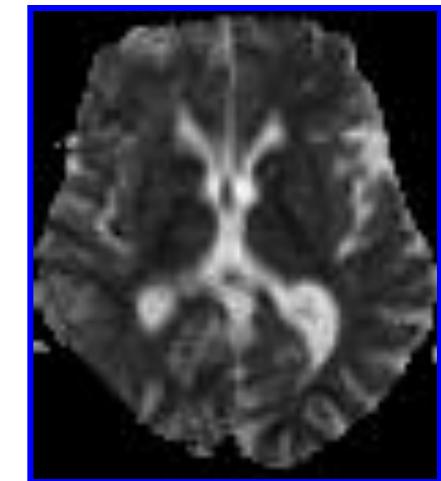
## Tensor model



- Principal diffusivities:  
diffusivities along the three principal directions



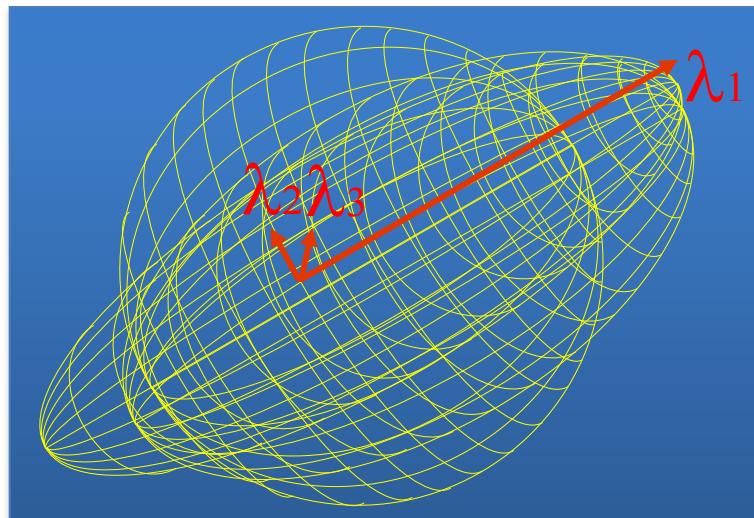
$$MD = (\lambda_1 + \lambda_2 + \lambda_3)/3$$



# DIFFUSION MRI

## Tensor model

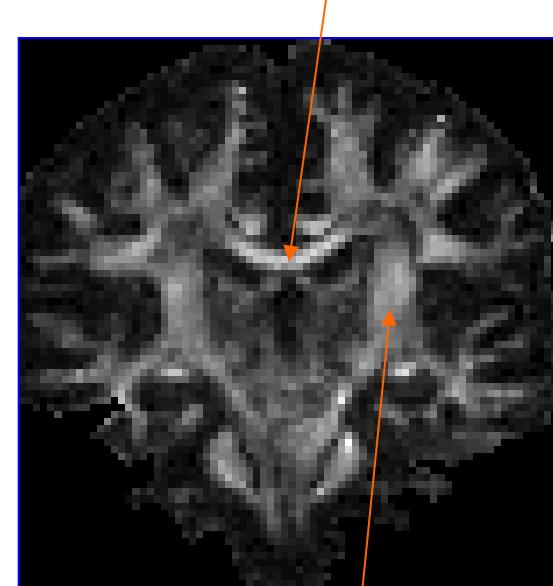
- Shape



- Fractional anisotropy index (FA):

$$FA = \sqrt{\frac{1}{2} \frac{\sqrt{(\lambda_1 - \lambda_2)^2 + (\lambda_2 - \lambda_3)^2 + (\lambda_3 - \lambda_1)^2}}{\sqrt{\lambda_1^2 + \lambda_2^2 + \lambda_3^2}}}$$

Corpus callosum



Internal capsule

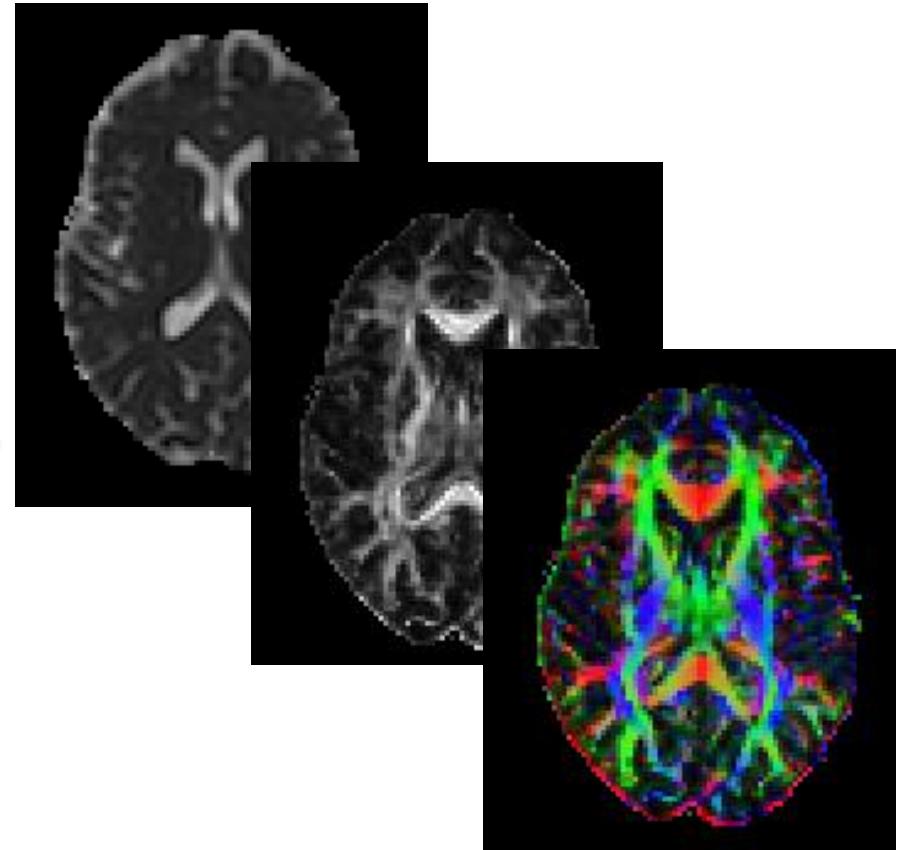
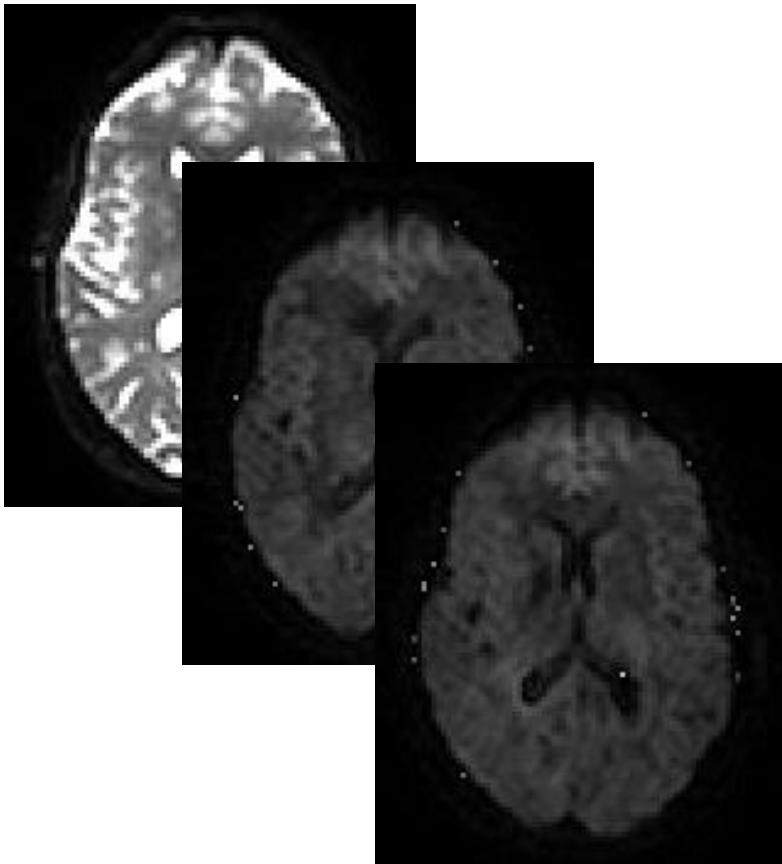
$$AxD = \lambda_1$$

$$RD = (\lambda_2 + \lambda_3)/2$$

# DIFFUSION MRI

## Diffusion tensor estimation

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# DIFFUSION MRI

## Diffusion techniques

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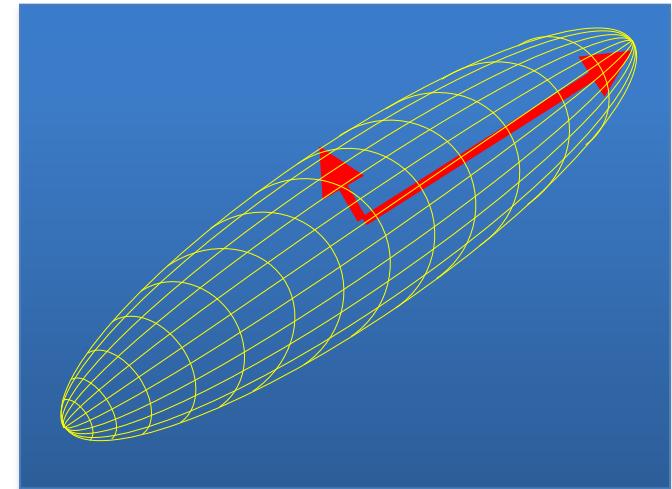
- ROI-based approaches
- Tractography-based approaches
- Voxel-wise approaches
  - Tract-based spatial statistics (TBSS)
  - Statistical parametric mapping (SPM)

# DIFFUSION MRI

## Tractography

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Assumption: only one direction is predominant in each voxel and diffusivity is greatest in this direction



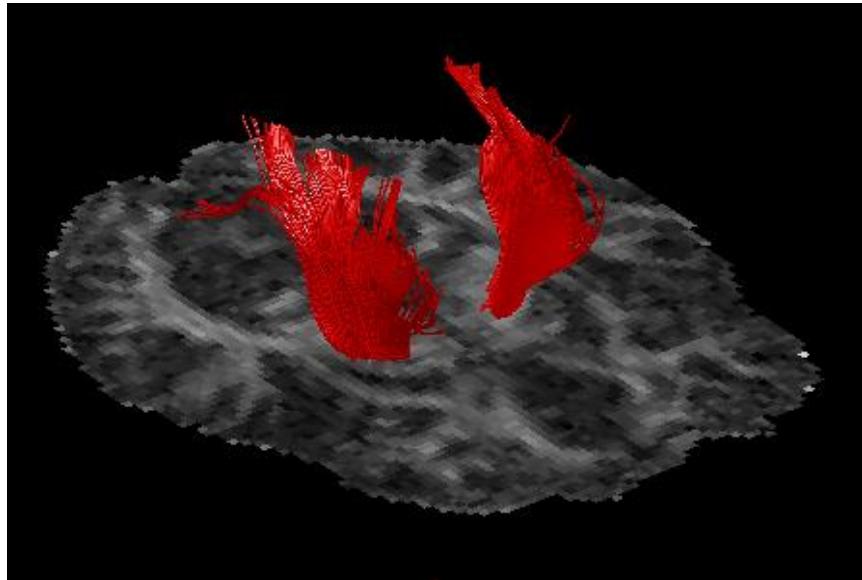
3D vector field representing fibre orientation in each voxel can be obtained through diagonalization of DT

Tracts are reconstructed by changing the fibre's direction as soon as a new pixel with a new information about direction is encountered.

# DIFFUSION MRI

## Tractography

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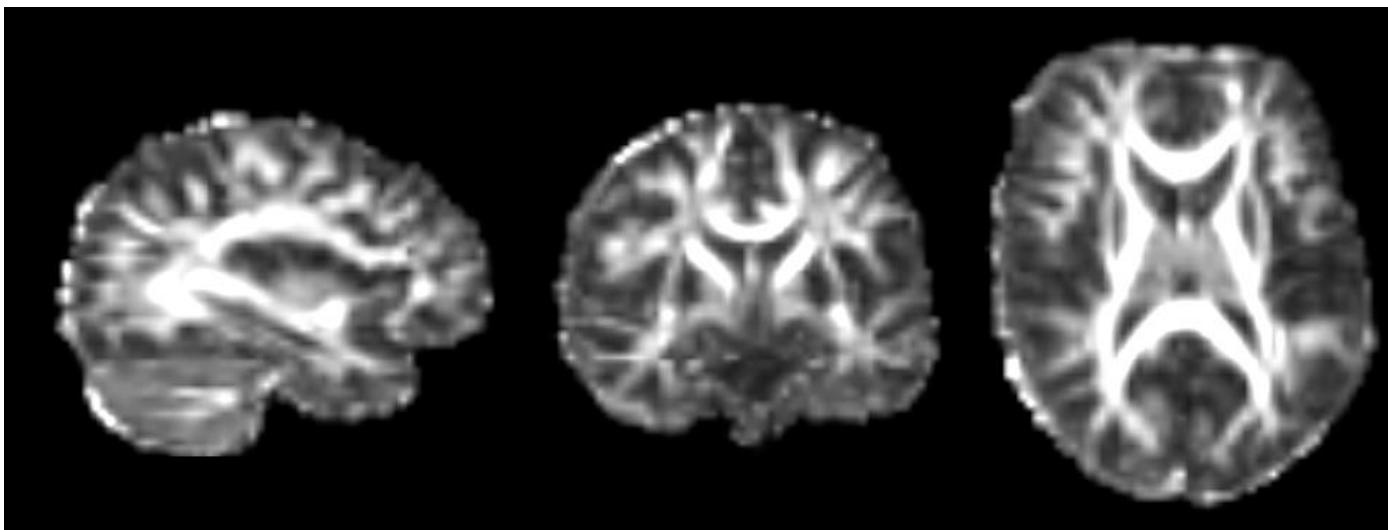


# DIFFUSION MRI

## TBSS

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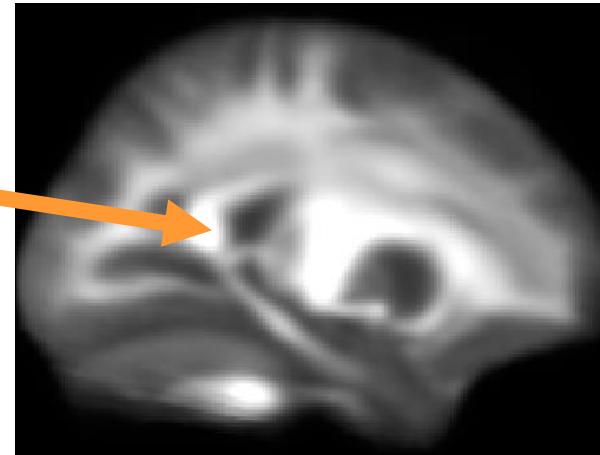
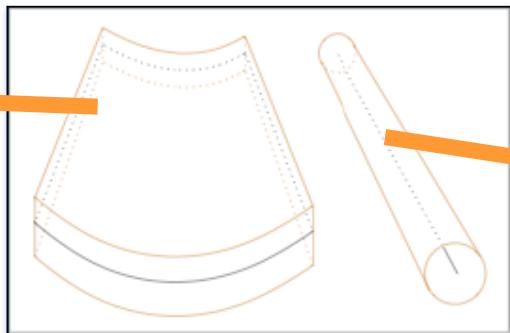
Use nonlinear registration to pre-align all subjects' FA  
(nonlinear reg: FNIRT)



[www.fmrib.ox.ac.uk/fsl/tbss/index.html](http://www.fmrib.ox.ac.uk/fsl/tbss/index.html)

# DIFFUSION MRI

## TBSS/Skeletonise

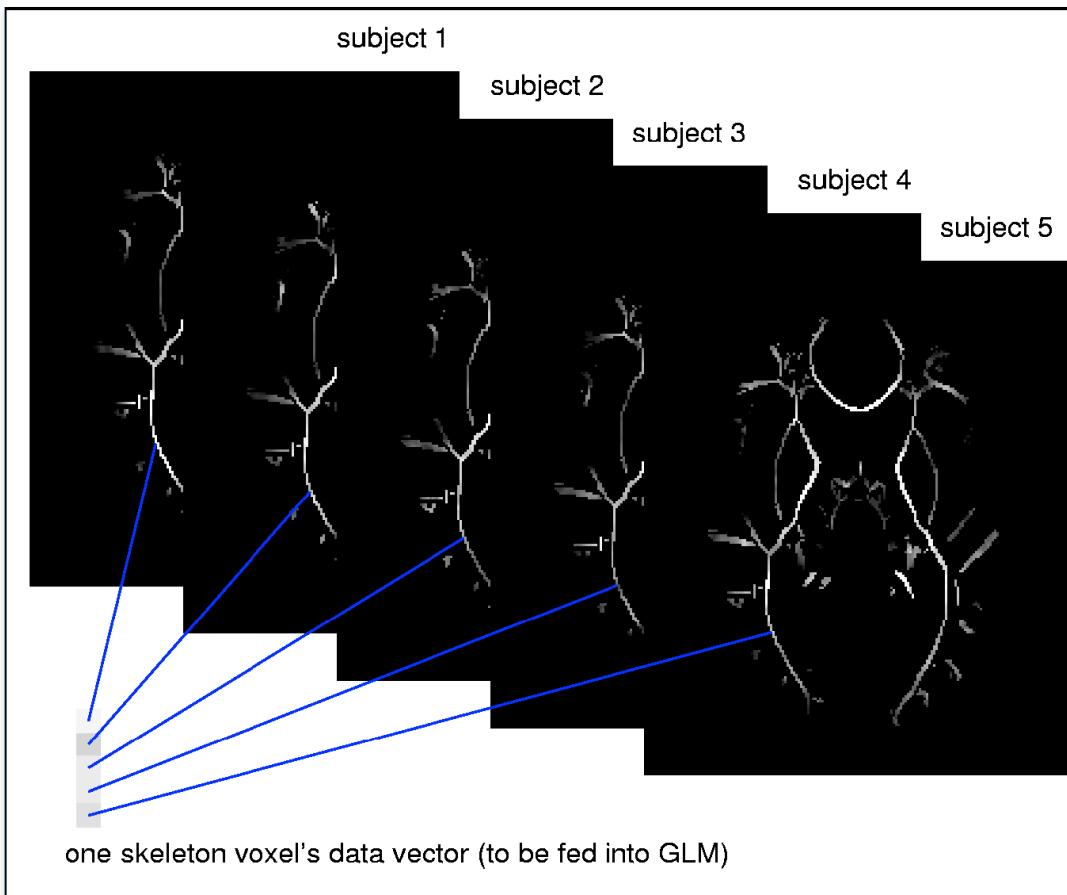


[www.fmrib.ox.ac.uk/fsl/tbss/index.html](http://www.fmrib.ox.ac.uk/fsl/tbss/index.html)

# DIFFUSION MRI

## TBSS

Do cross-subject voxelwise stats on skeleton-projected FA Threshold, (e.g., permutation testing, including multiple comparison correction)



# DIFFUSION MRI

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# DIFFUSION MRI

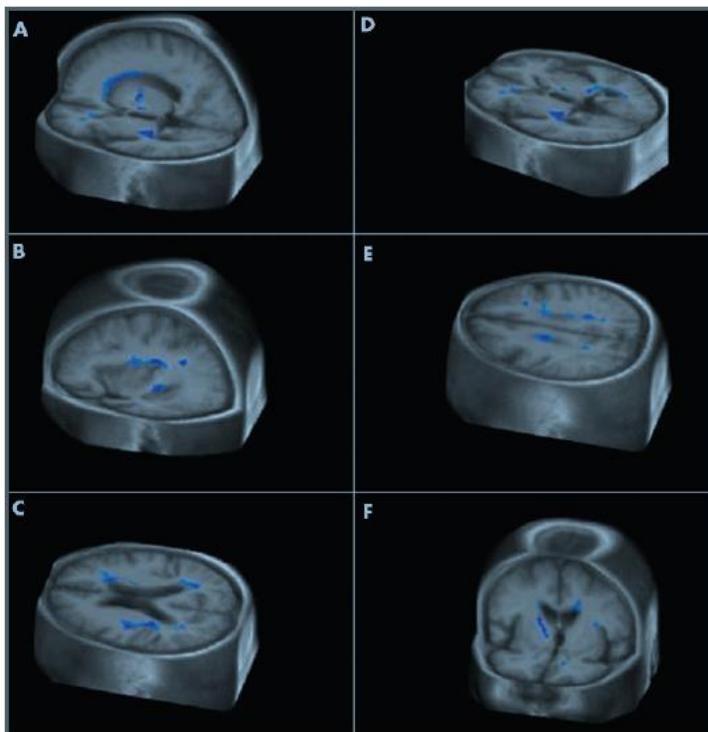
## Outline of the presentation

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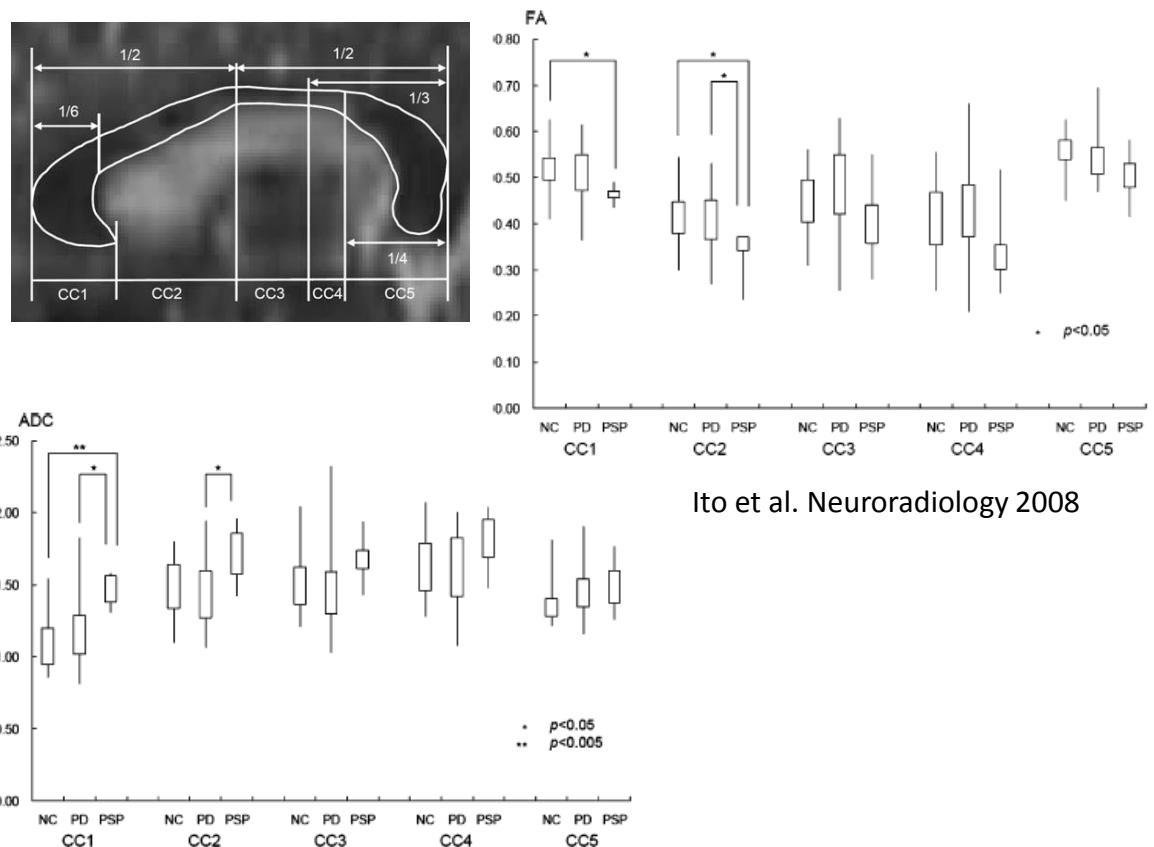
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# DIFFUSION MRI

## PSP

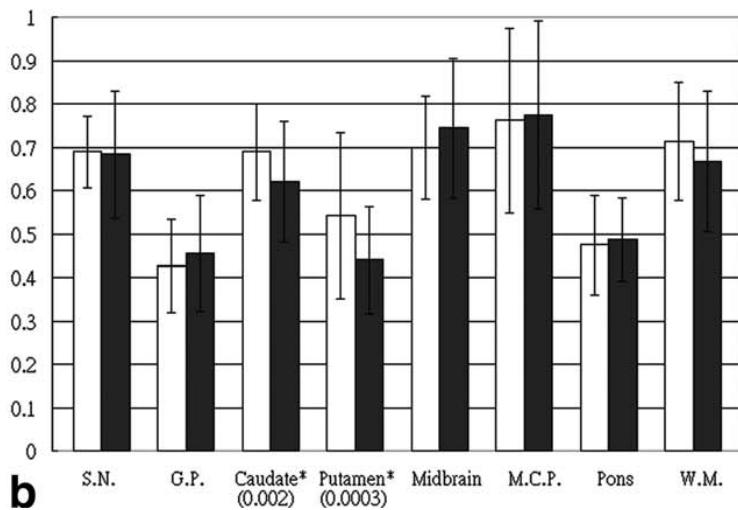
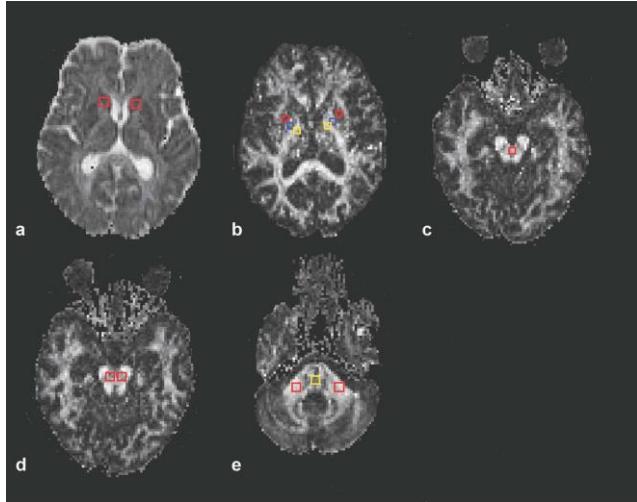


Padovani et al. JNNP 2006

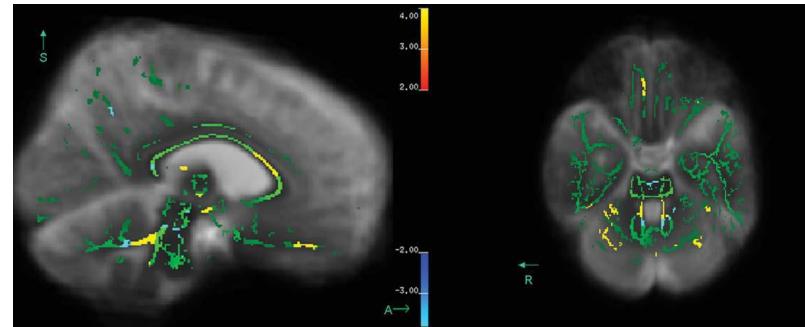


# DIFFUSION MRI

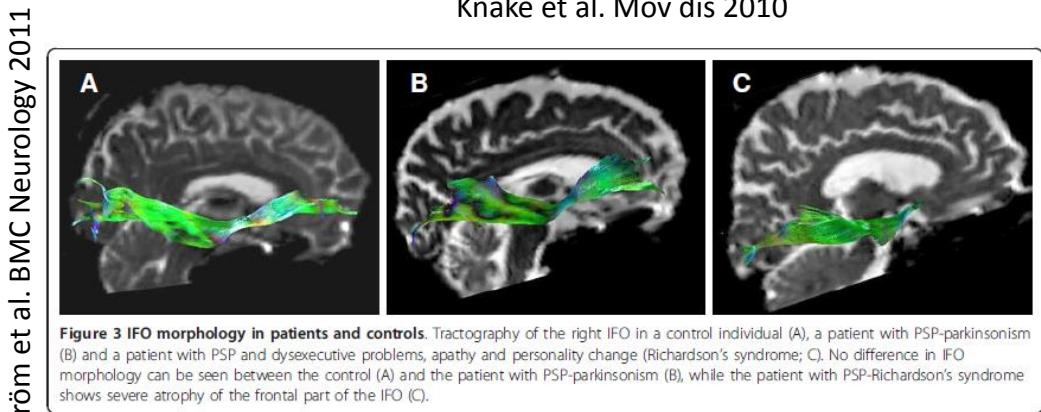
## PSP



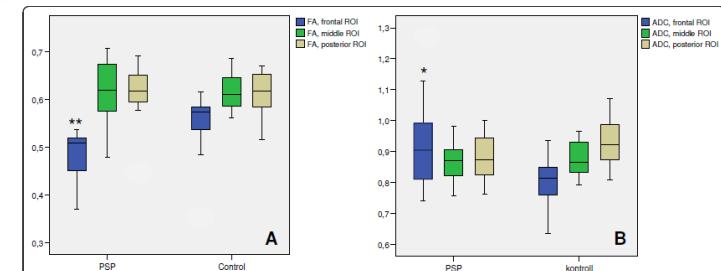
Wang et al. JMRI 2010



Knake et al. Mov dis 2010



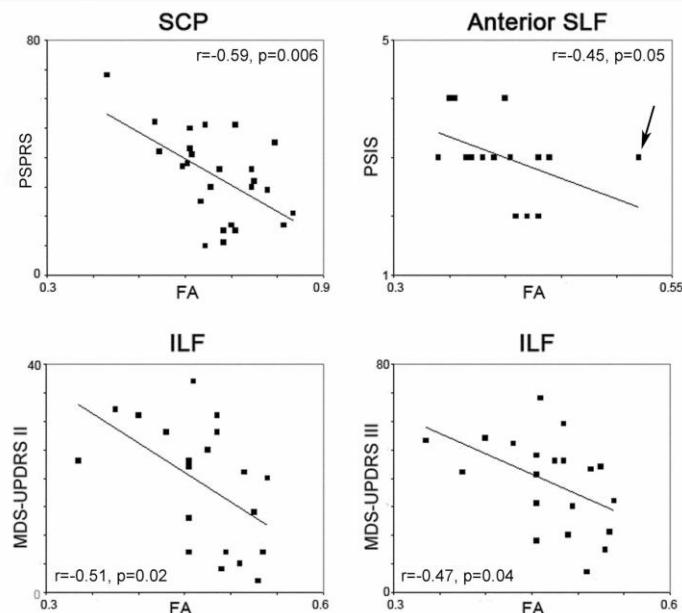
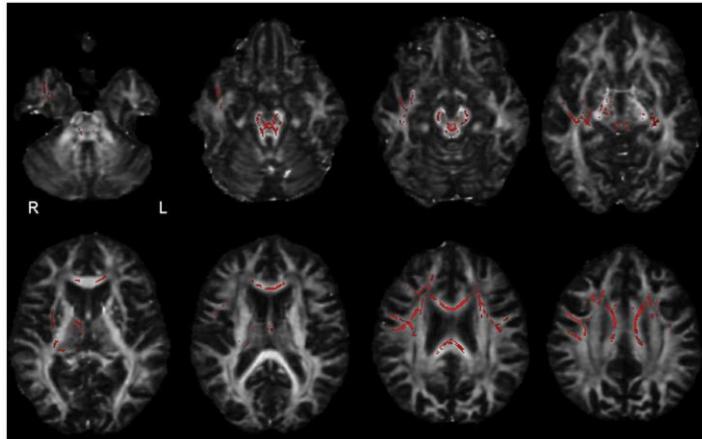
**Figure 3** IFO morphology in patients and controls. Tractography of the right IFO in a control individual (A), a patient with PSP-parkinsonism (B) and a patient with PSP and dysexecutive problems, apathy and personality change (Richardson's syndrome; C). No difference in IFO morphology can be seen between the control (A) and the patient with PSP-parkinsonism (B), while the patient with PSP-Richardson's syndrome shows severe atrophy of the frontal part of the IFO (C).



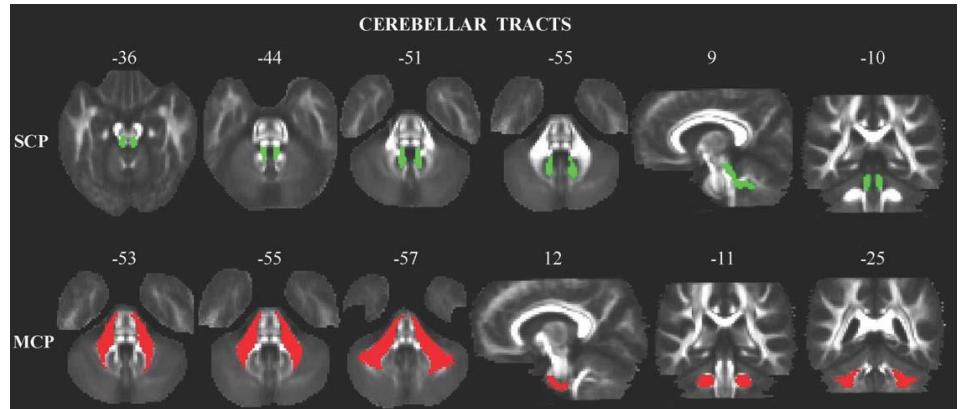
**Figure 4** FA and ADC in the IFO. Box-plot of FA (A) and ADC (B,  $\text{mm}^2/\text{s}$ ) values in the frontal, middle and posterior ROIs of the IFO (see text for description of location). In the frontal ROI the FA value was significantly decreased (\*\*;  $p < 0.01$ ), while the ADC value was significantly increased (\*;  $p < 0.05$ ), compared to controls. No difference in either FA or ADC was seen between controls and patients with PSP for the middle or posterior ROIs.

# DIFFUSION MRI

## PSP

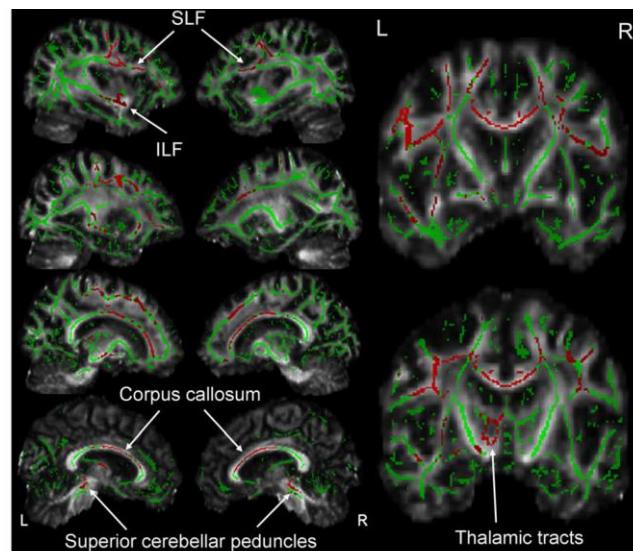


Whitwell et al. Arch Neurol. 2011



Selective damage of SCP

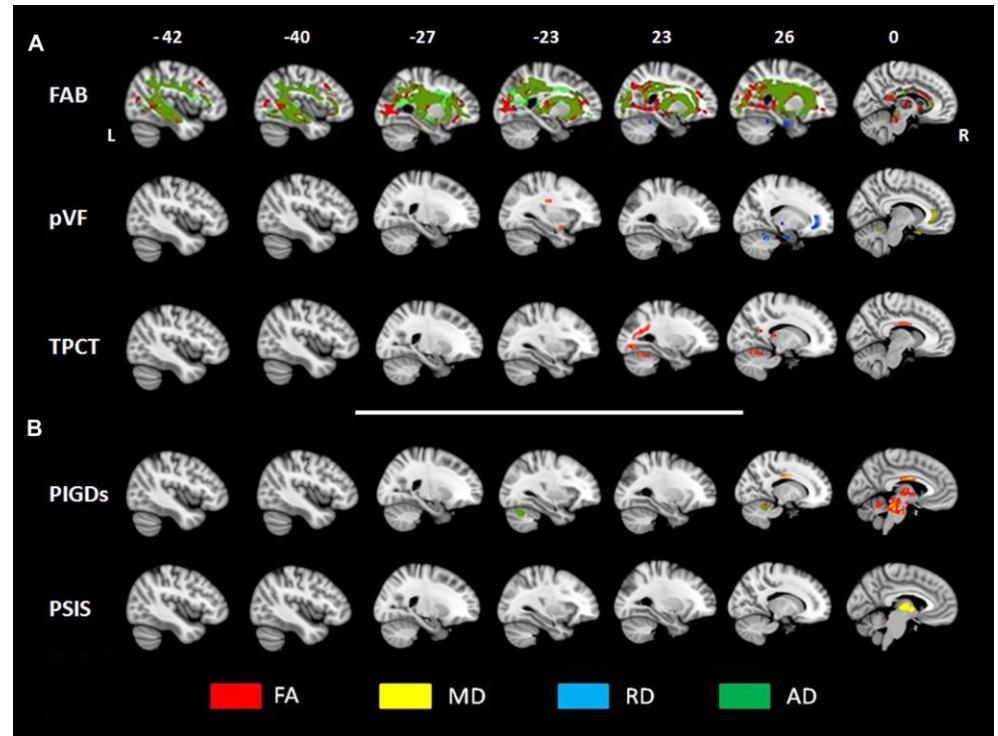
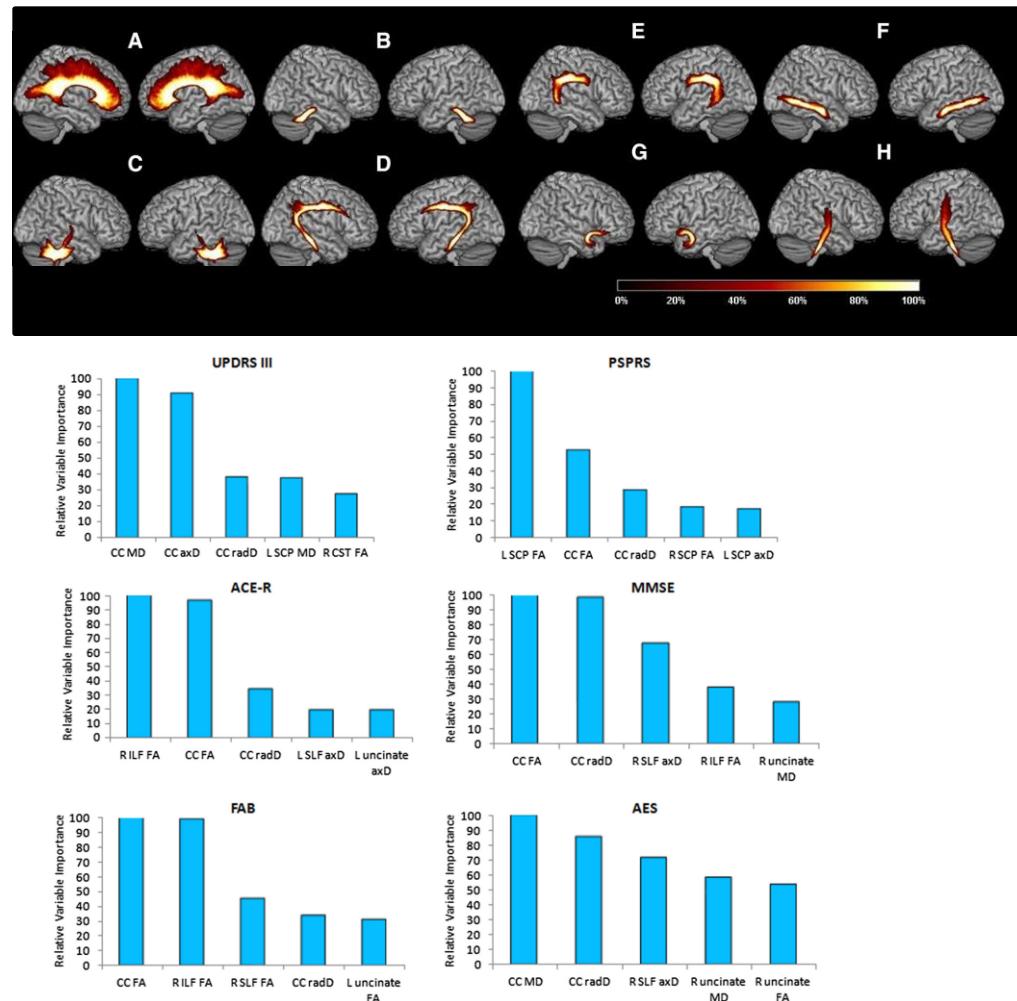
Canu et al. Mov Dis. 2011



Whitwell et al. Parkinsonism Relat Disord. 2011

# DIFFUSION MRI

## PSP/Correlations



Tessitore et al Neurobiol Aging. 2014

# DIFFUSION MRI

## Outline of the presentation

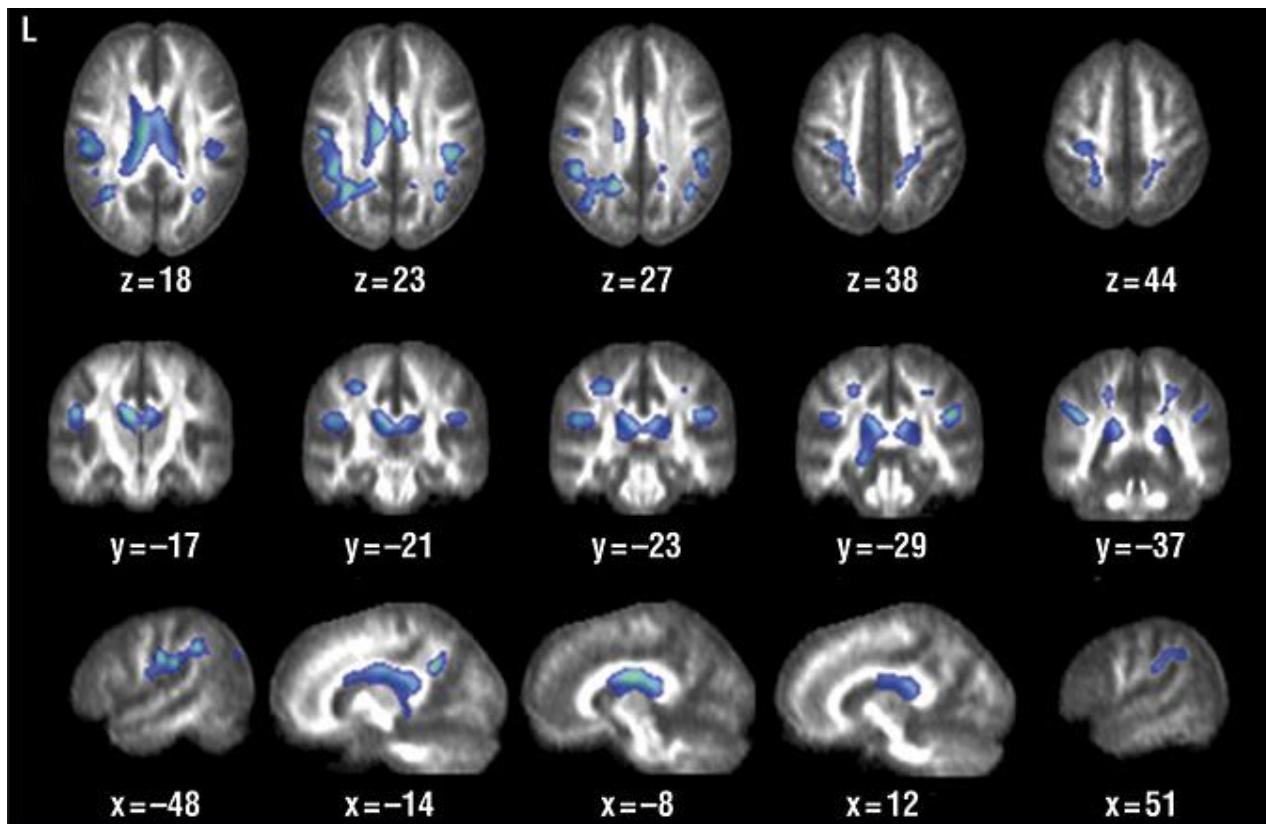
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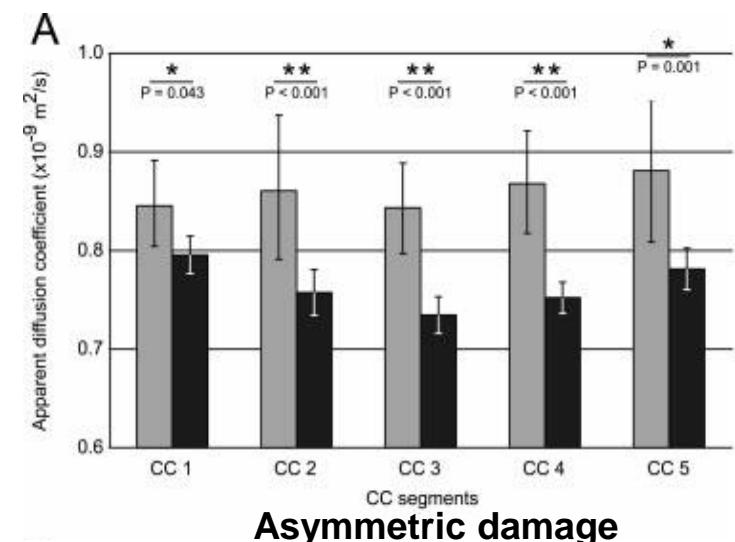
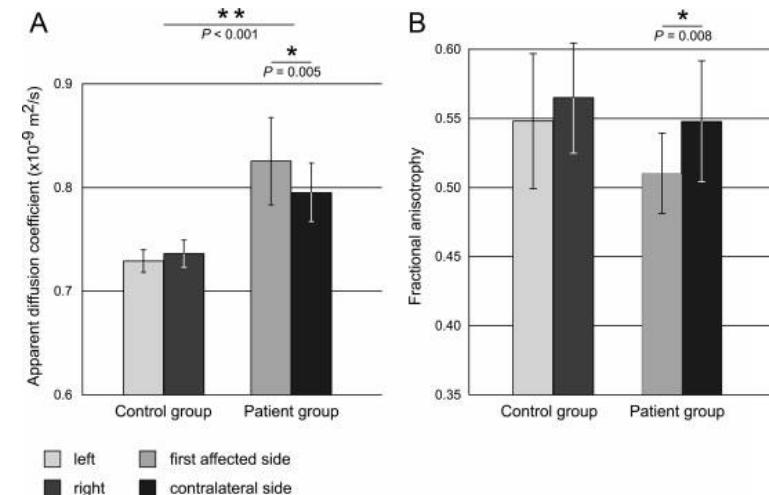
## CBS

### FA: CBS patients vs. controls



Corpus callosum, fronto-parietal connections

Borroni et al., Arch Neurol. 2008



Boelmans et al. Mov Dis. 2009

# DIFFUSION MRI

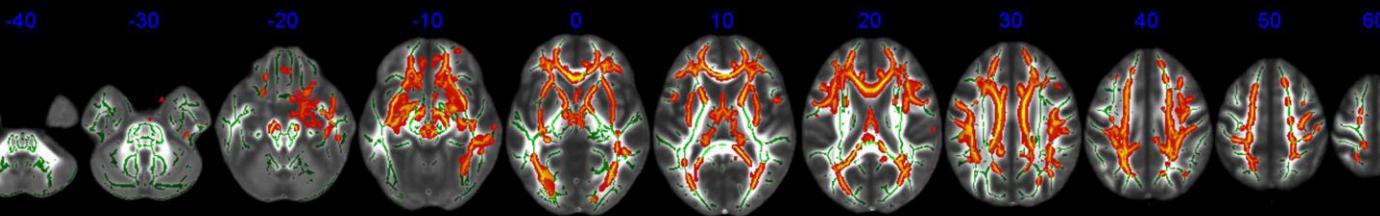
## CBS

DTI metrics changes in CBS

A

### FRACTIONAL ANISOTROPY

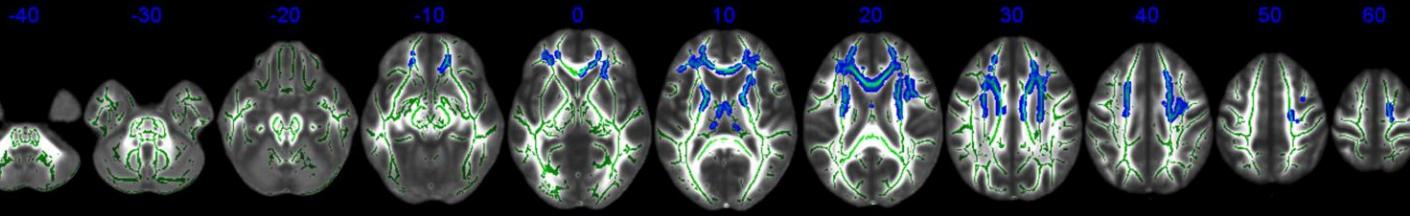
CTL > CBS



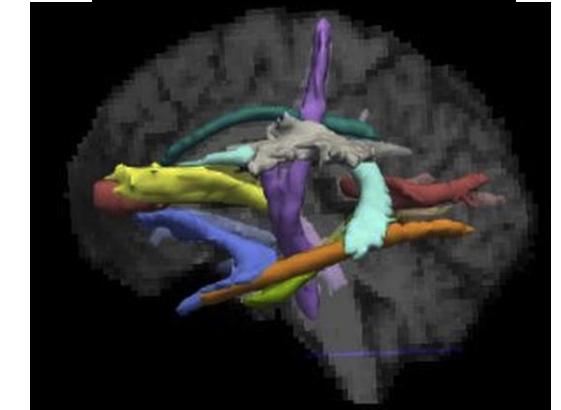
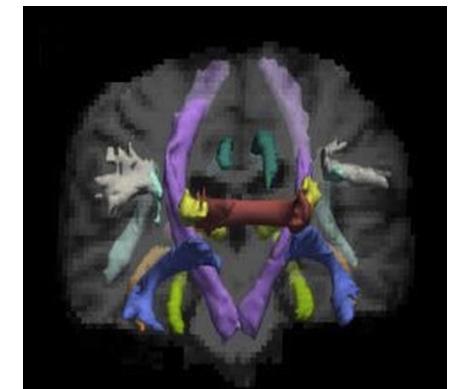
B

### MEAN DIFFUSIVITY

CTL < CBS



Most tracts are altered in CBS



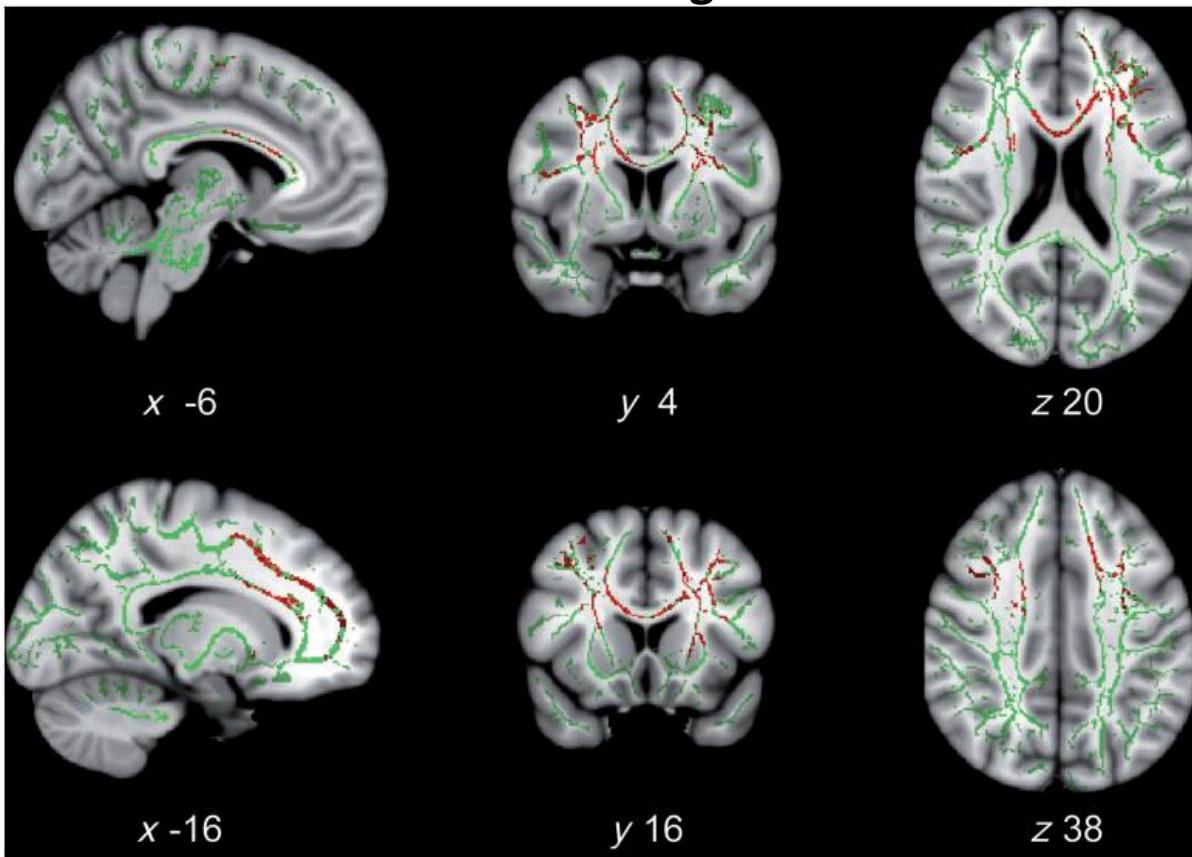
Tovar-Moll et al. Plos One. 2014

Upadhyay et al. Neurobiol Aging. 2016

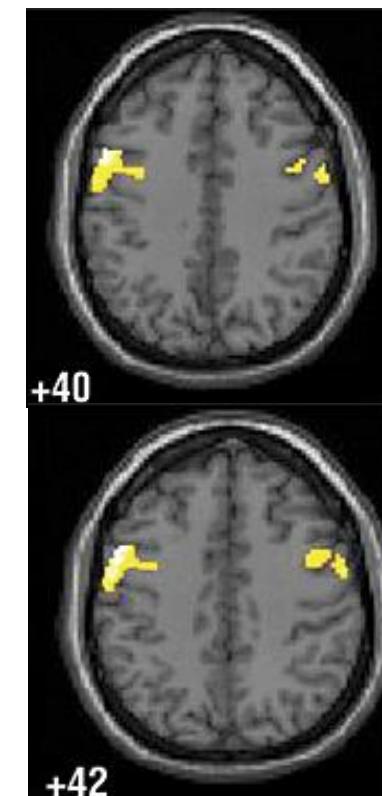
# DIFFUSION MRI

## CBS/Correlations

MD correlates to action binding in CBS



Sensorimotor WM FA decrease  
vs. limb apraxia



# DIFFUSION MRI

## Outline of the presentation

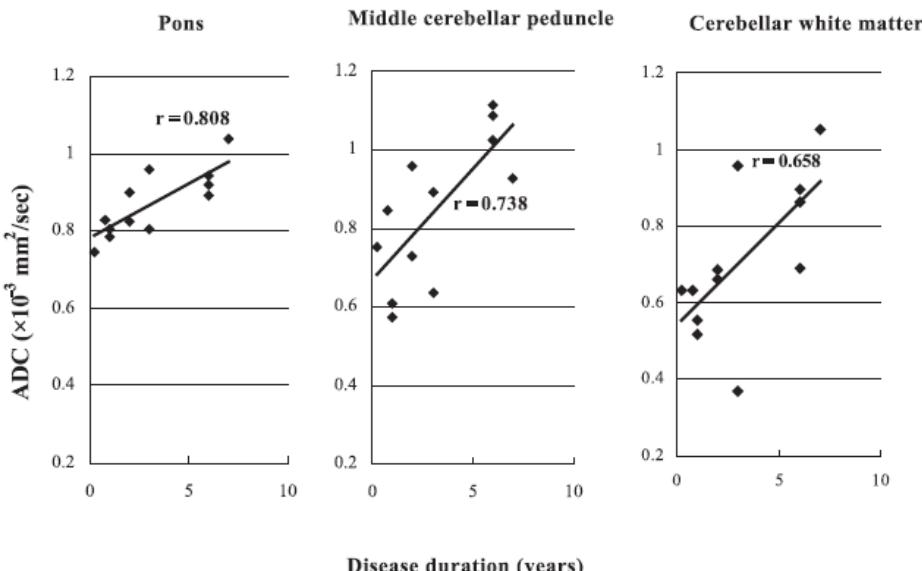
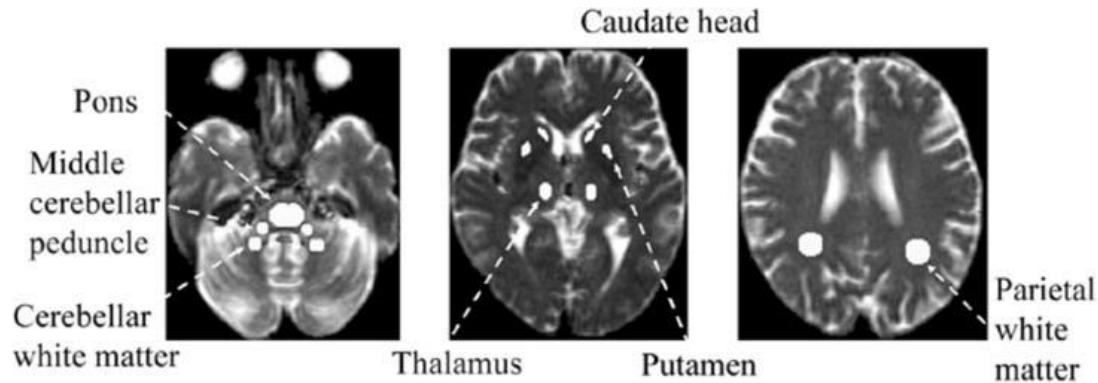
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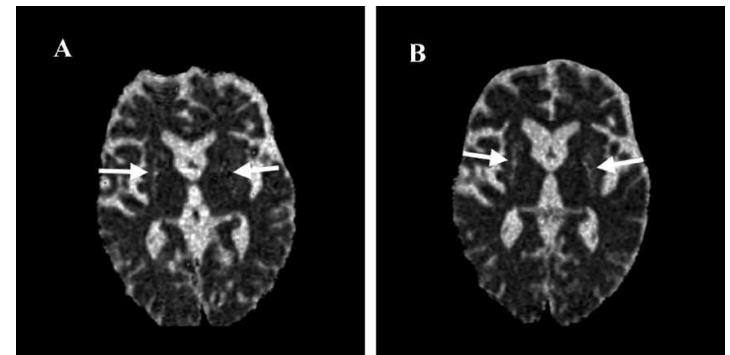
## MSA

Altered ADC in the pons and MCP of MSA-C patients

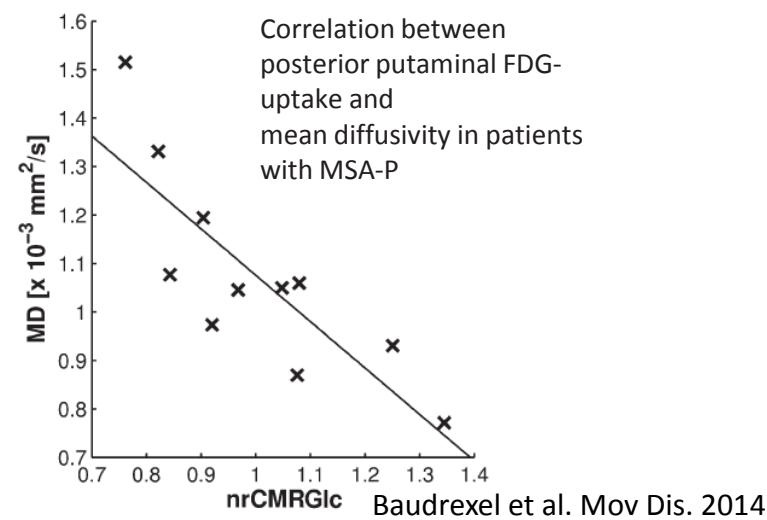


Kanazawa et al. J Neurol. 2004

Putaminal degeneration in MSA-P

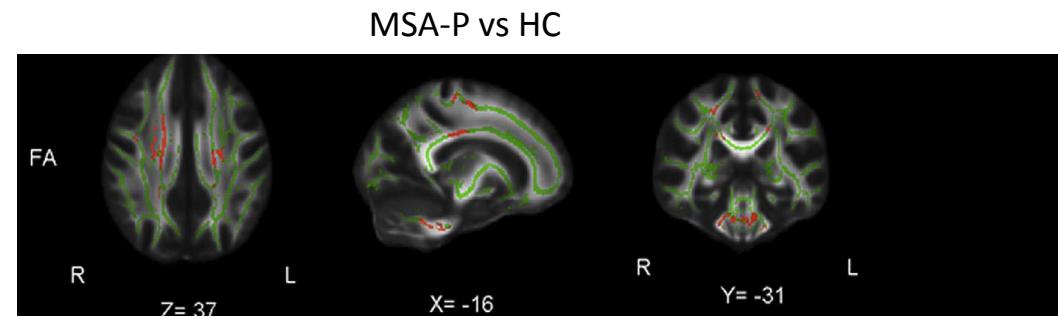
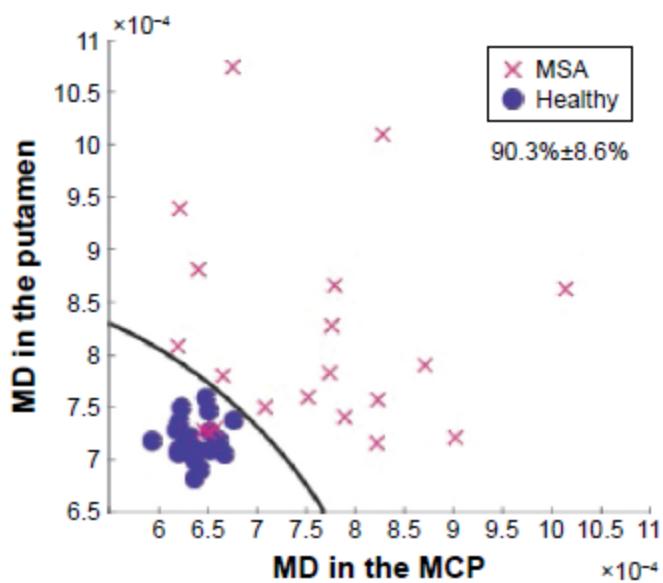
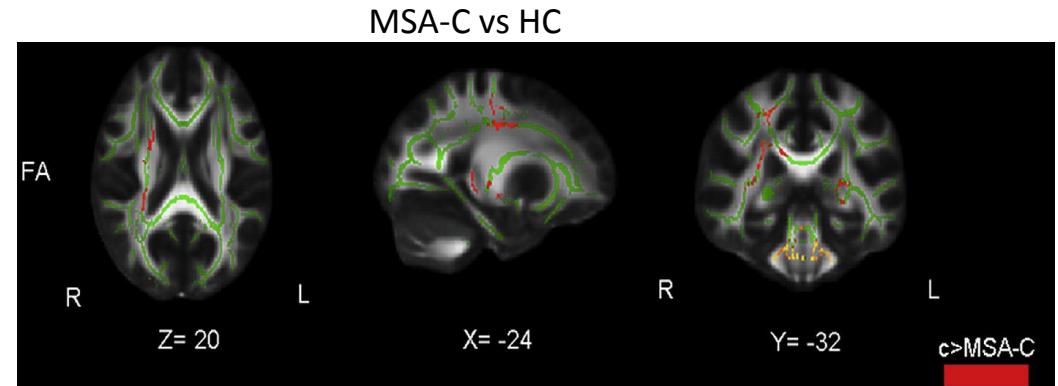
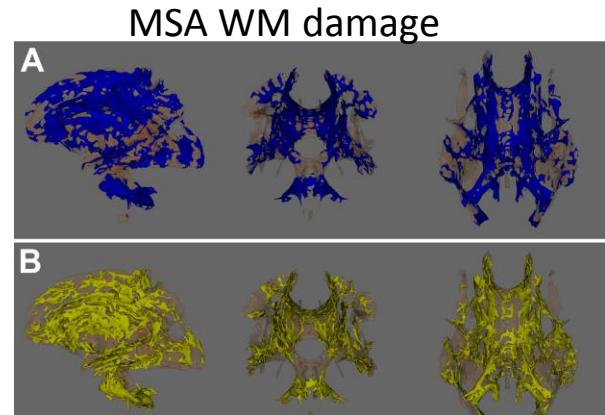


Seppi et al. Neuroimage 2006



# DIFFUSION MRI

## MSA



Ji et al. Parkinsonism Relat Disord. 2014

# DIFFUSION MRI

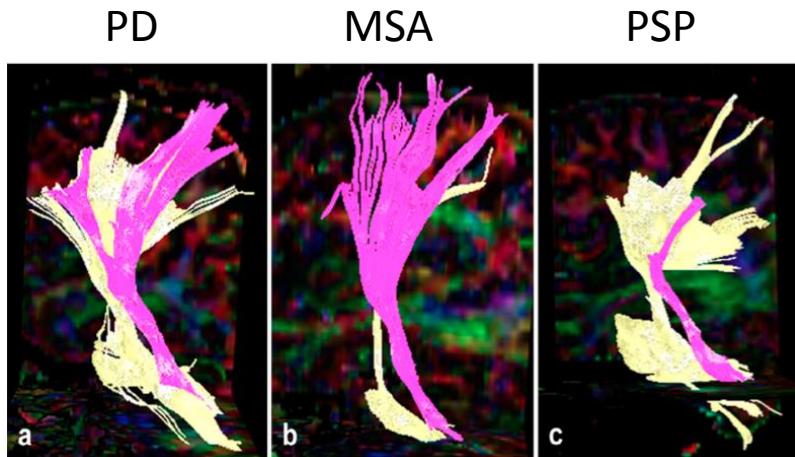
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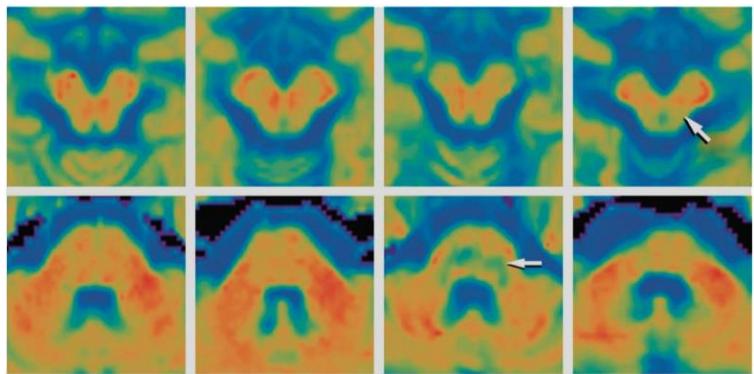
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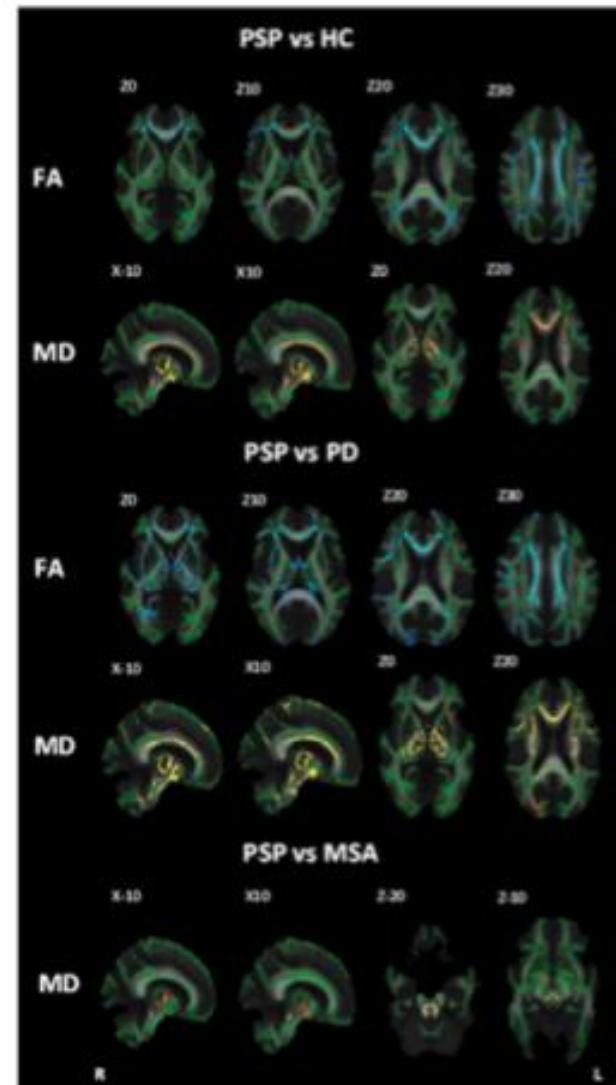
## Differential diagnosis



Nilsson et al. Neuroradiology. 2007



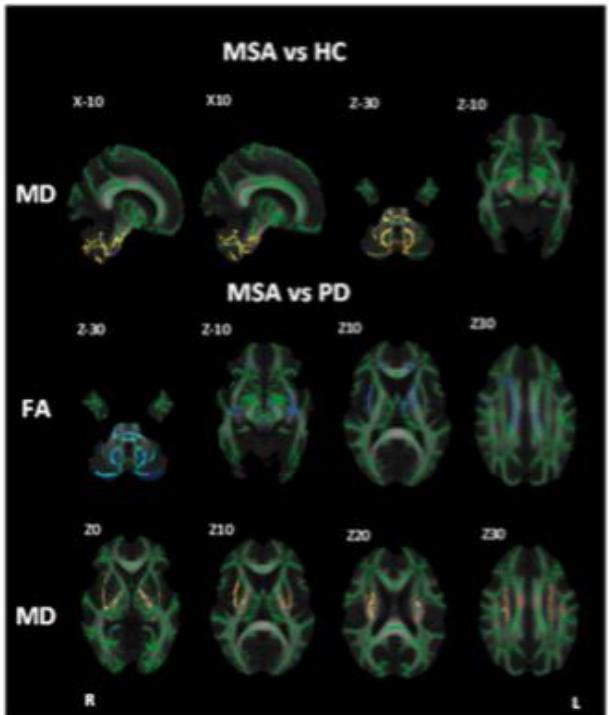
Ito et al. Neuroreport. 2015



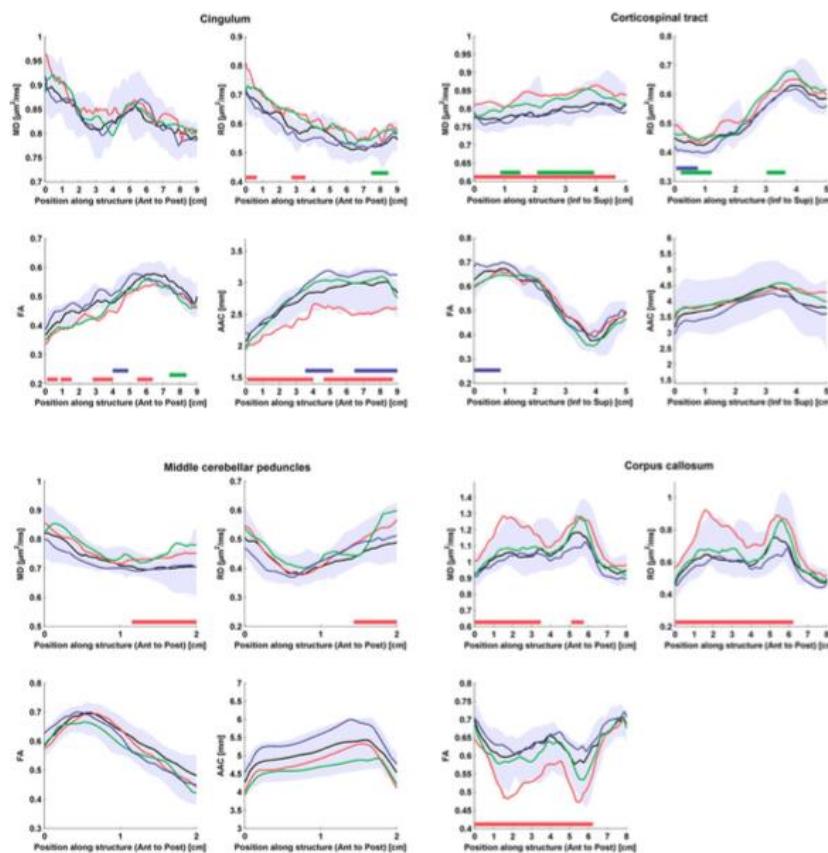
Worker et al. Plos one. 2014

# DIFFUSION MRI

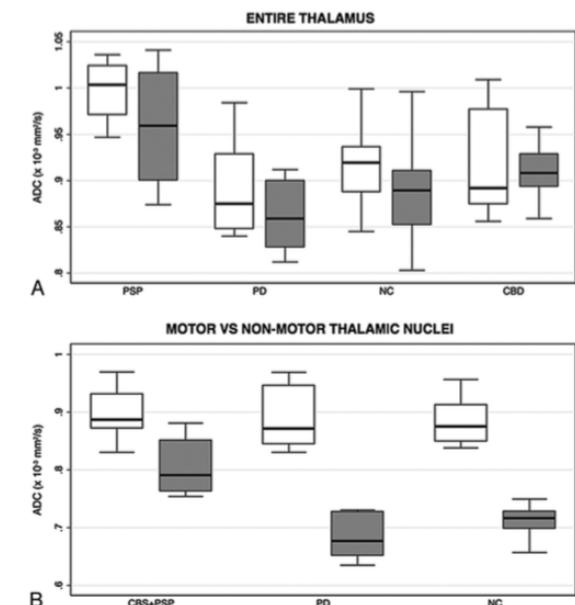
## Differential diagnosis



Worker et al. Plos one. 2014



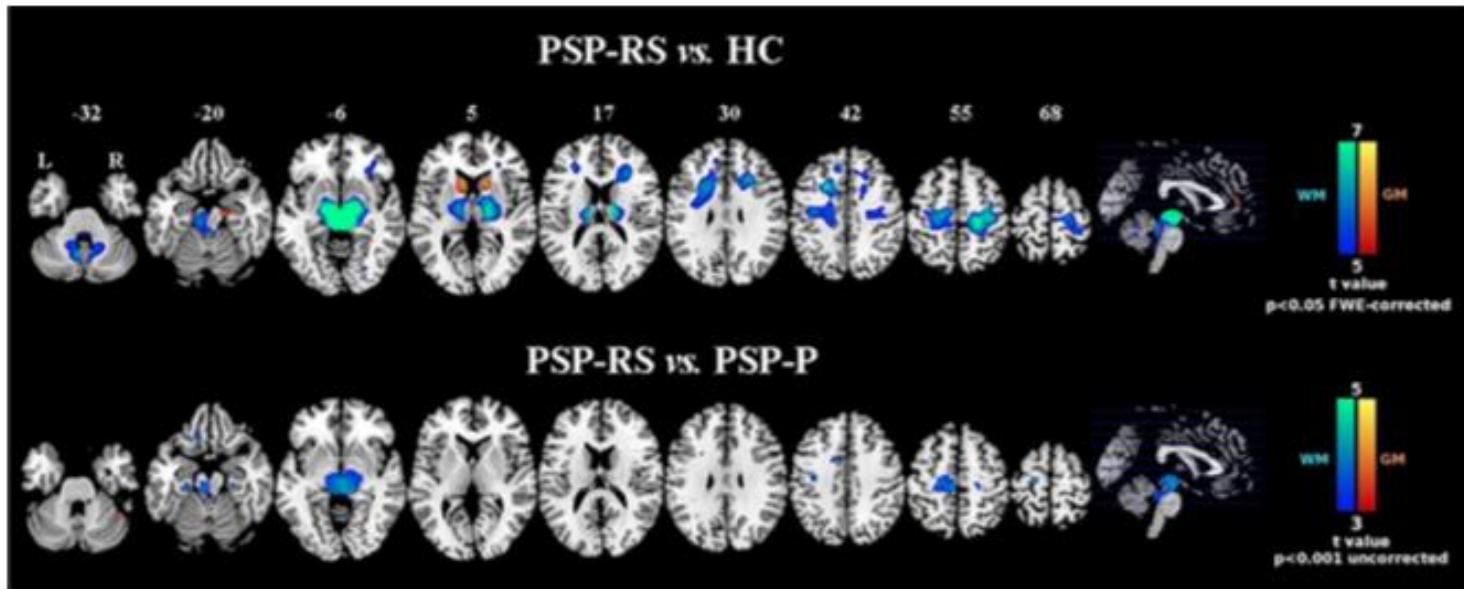
Surova et al. Plos one. 2013



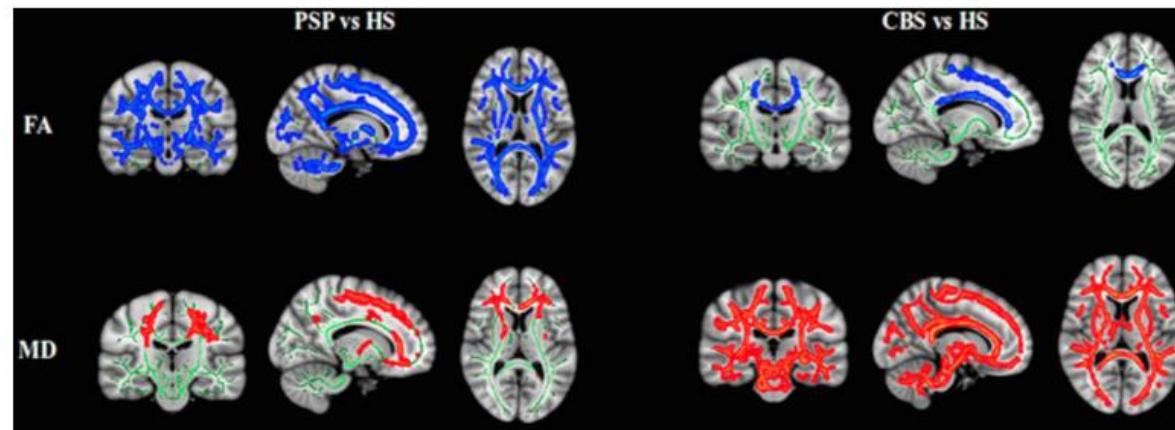
Hess et al. Brain. 2014

# DIFFUSION MRI

## Differential diagnosis



Agosta et al. Neurobiol Aging. 2012

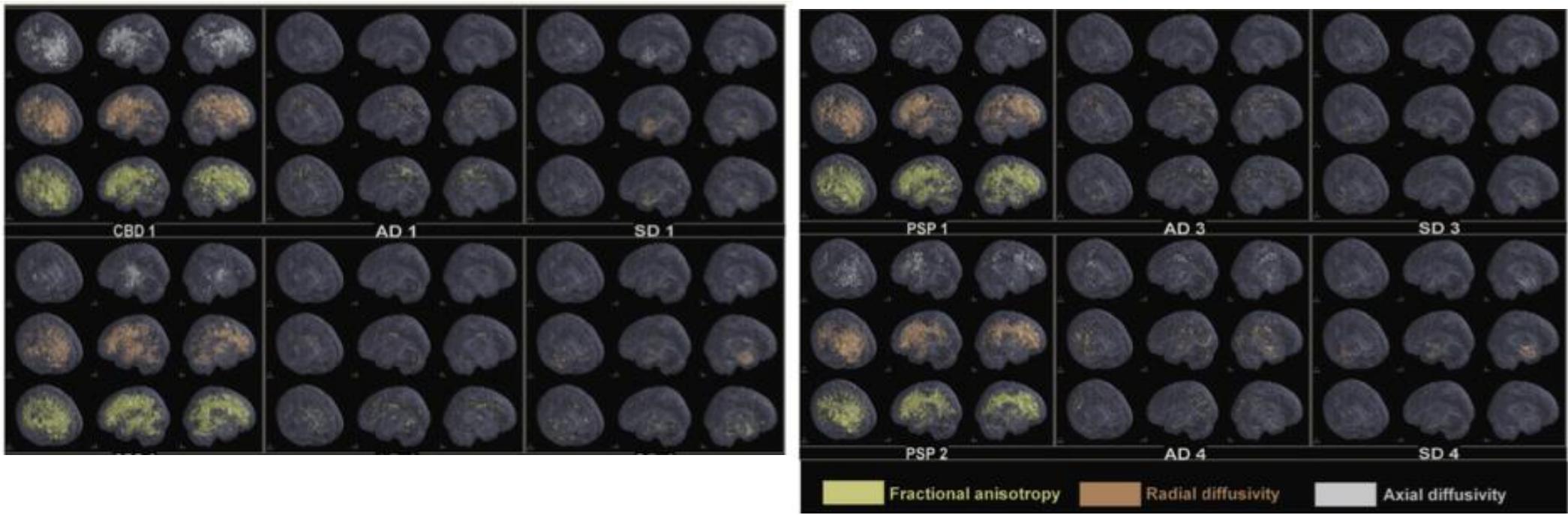


Upadhyay et al. J Neurol. 2016

# DIFFUSION MRI

## Differential diagnosis/Single subject

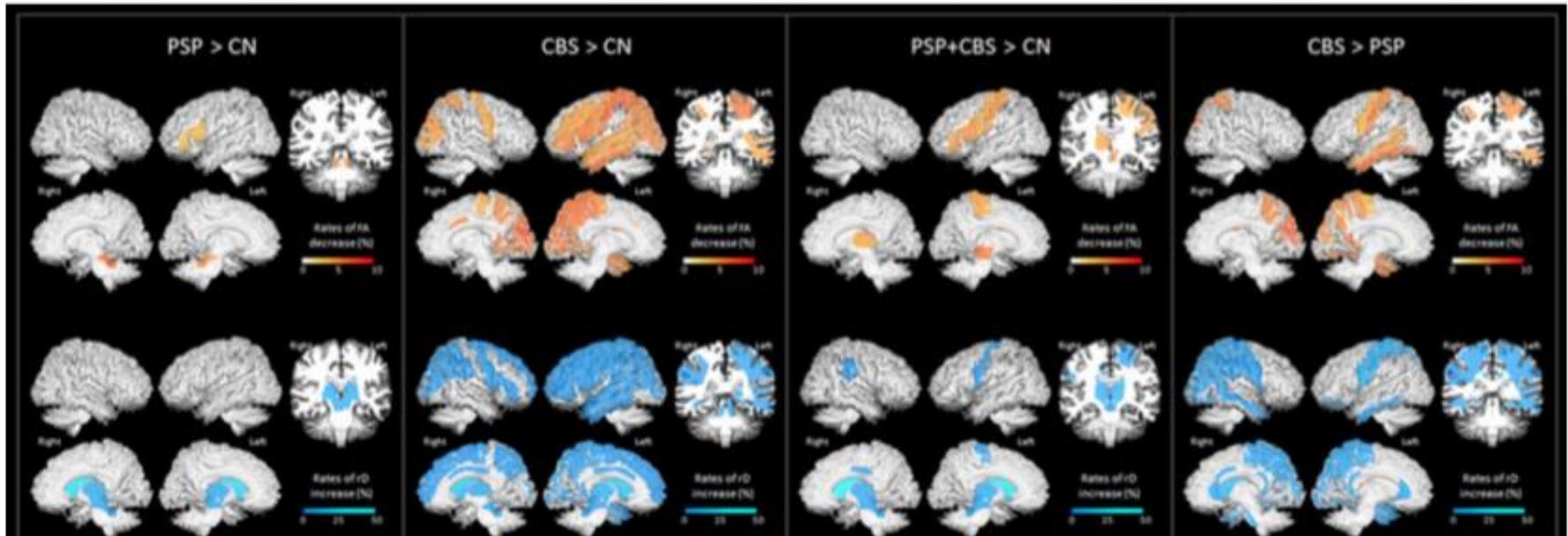
Single subject diagnosis of PSP or CBD



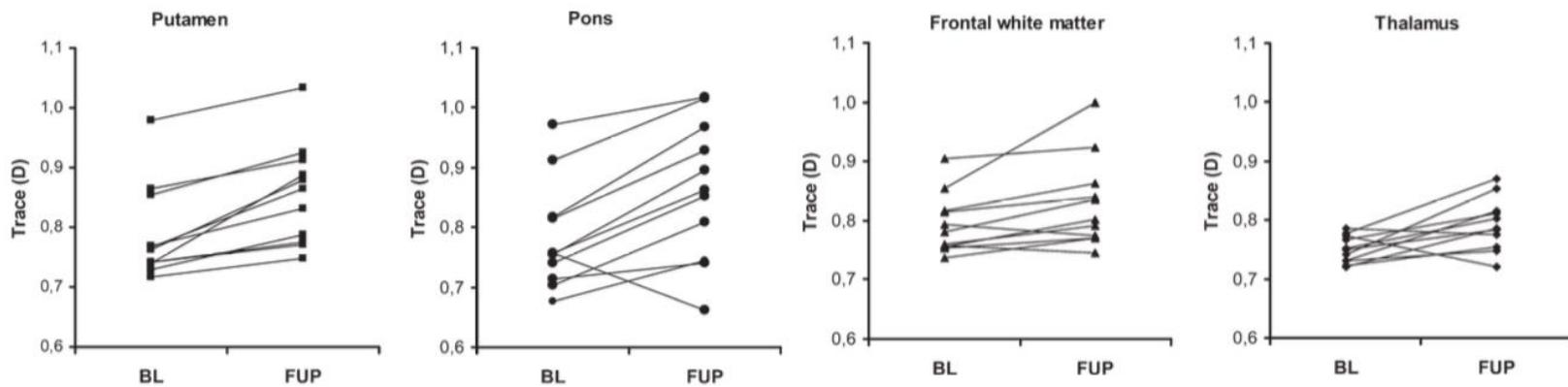
# DIFFUSION MRI

## Monitoring disease progression

Zhang et al. Plos one. 2016



Pellecchia et al. Mov Dis. 2011





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# Neuroimaging Research Unit & Neurodegenerative Diseases Group

**Director: M. Filippi**

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