



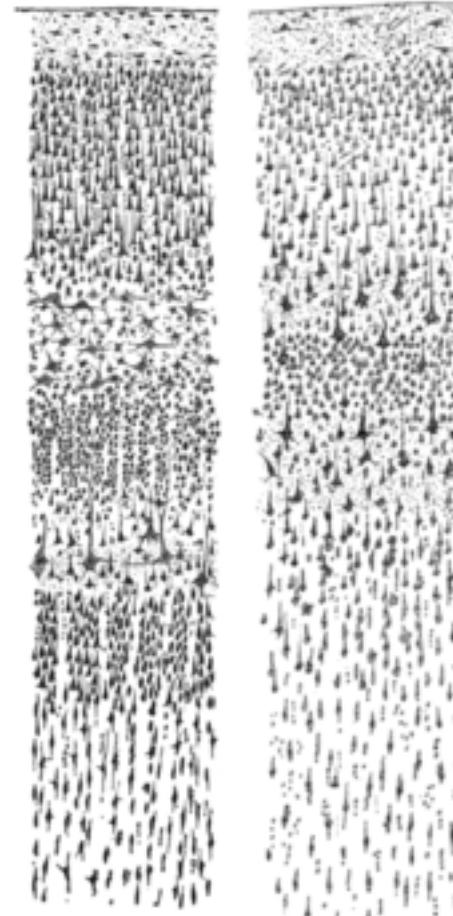
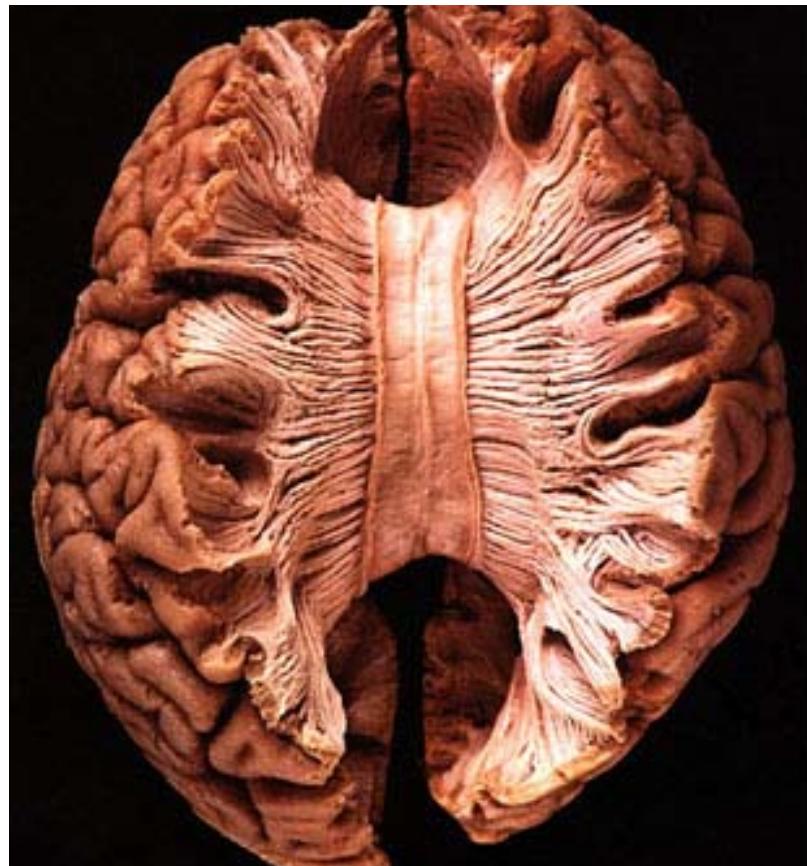
## Detailed laminar characteristics of the human neocortex revealed by NODDI and histology

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University College London, UK ..... Hui Zhang  
Max Planck Institute, Cologne, D ..... Dirk Wiedermann  
UMC St. Radboud, Nijmegen, NL ..... Benno Küsters  
Donders Institute, Nijmegen, NL ..... David Norris  
UMC St. Radboud, Nijmegen, NL ..... Anne-Marie van Cappellen van Walsum





## Neocortex and cytoarchitectonic mapping



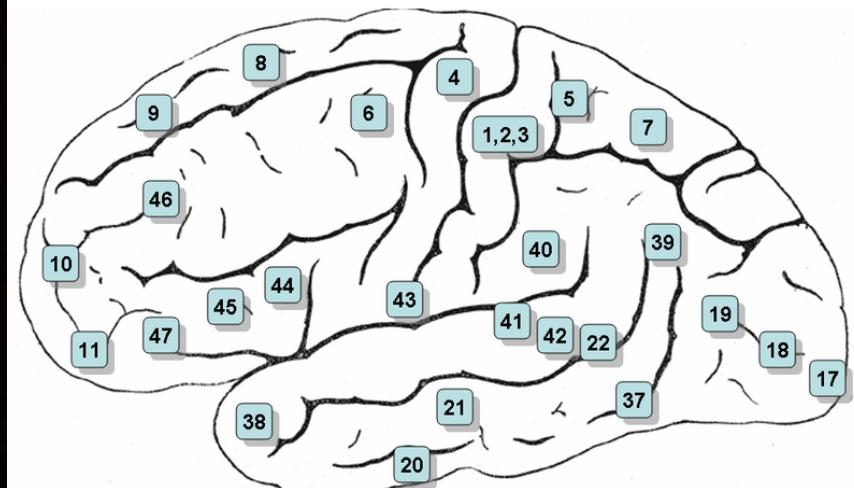
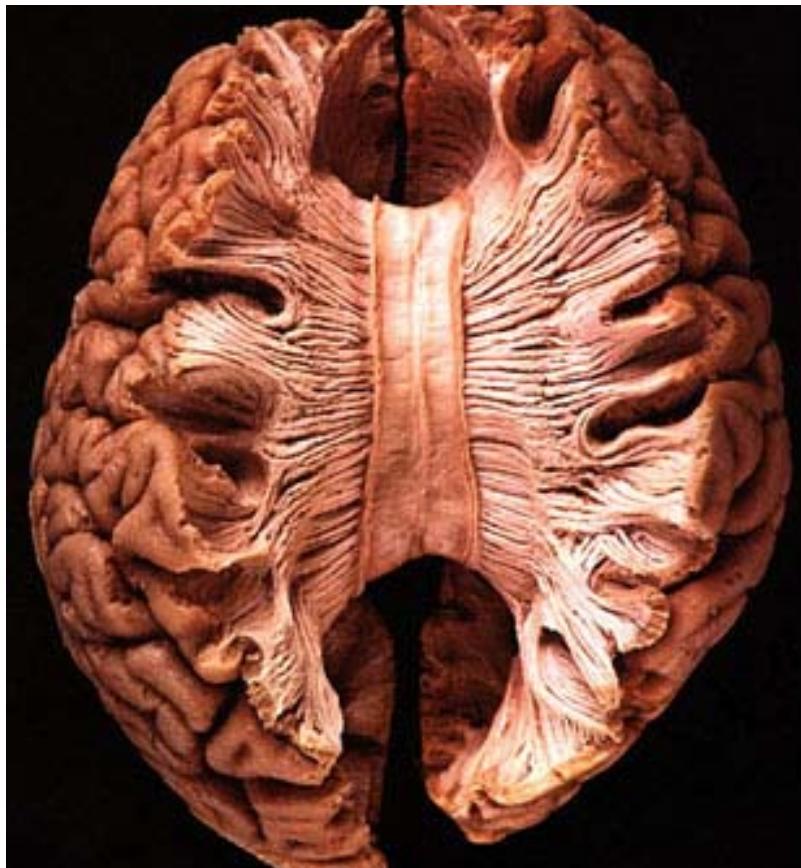
Visual  
cortex

Motor  
cortex





## Neocortex and cytoarchitectonic mapping



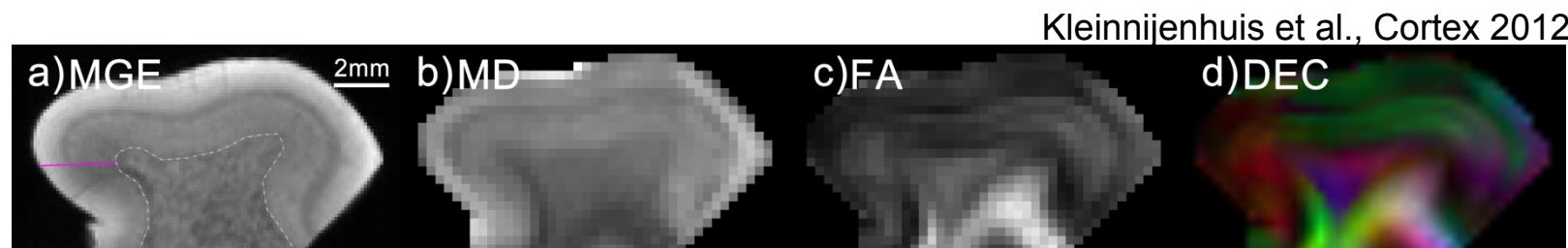
Broadmann areas





## Towards in vivo cytoarchitecture

- Diffusion can be used as structural probe
- Tensor metrics vary over cortical layers

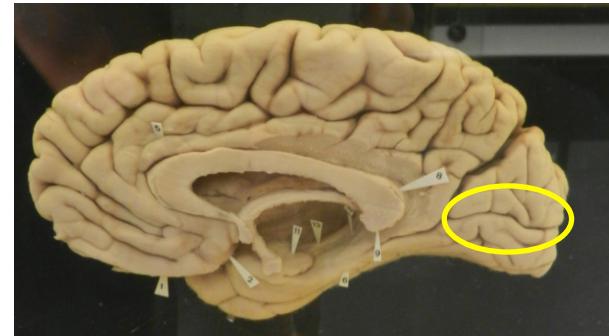


- Purpose of the current work:
  - extension to multishell
  - histology



## Samples of human V1

- post-mortem interval (< 24 h)
- fixed in formalin (> 1 month)
- 1 cm<sup>3</sup> calcarine sulcus (V1)
- soaked in phosphate buffered saline (> 72 h)
- scanned in proton-free liquid





## Diffusion Weighted Imaging

- System:
  - 9.4T Bruker BioSpec;  $G_{\max} = 660 \text{ mT/m}$
  - cryogenic mouse brain coil (20-30 K)
- PGSE with segmented EPI readout
- TR/TE = 6750/26 ms
- 0.2 mm isotropic voxels
- 8 shells x 384 (sample A) / 54 directions (sample B)
  - $b = [0 \ 1000 \ 3000 \ 4000 \ 8000 \ 12000 \ 16000 \ 20000] \text{ s mm}^{-2}$
  - $\delta/\Delta = 8/12 \text{ ms}$

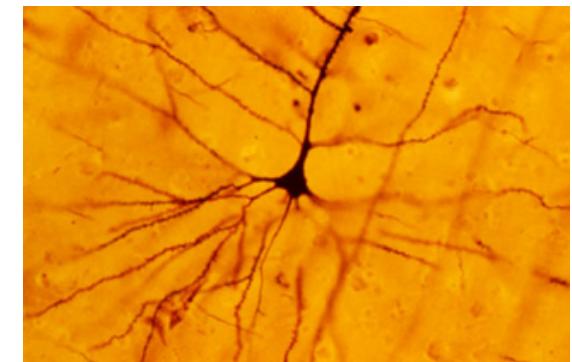




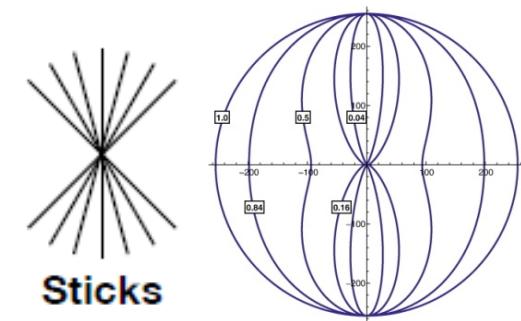
## Neurite Orientation Dispersion and Density Imaging

- NODDI multicompartment tissue model (Zhang et al., NI 2012)

1. neurite volume fraction
  2. extra-cellular volume fraction
  3. isotropic volume fraction
  4. isotropic restriction compartment
- ➔ ex vivo only (Alexander et al, 2010)

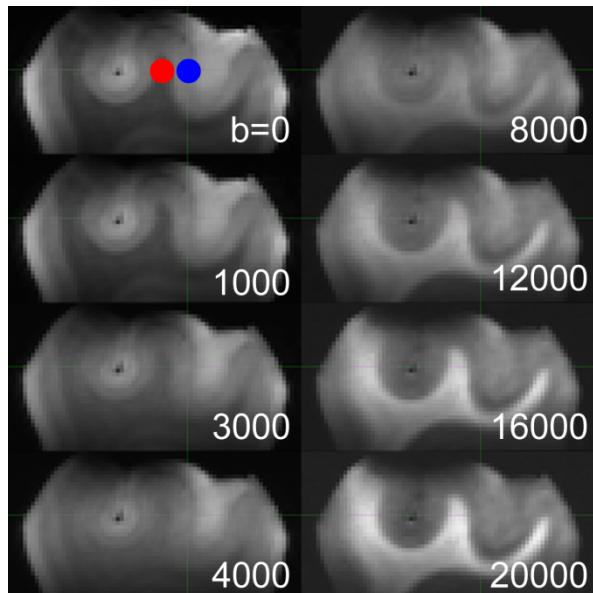


- Watson distribution
  - ➔ mean orientation  $\mu$  and concentration  $\kappa$
  - ➔ modeling WM & GM

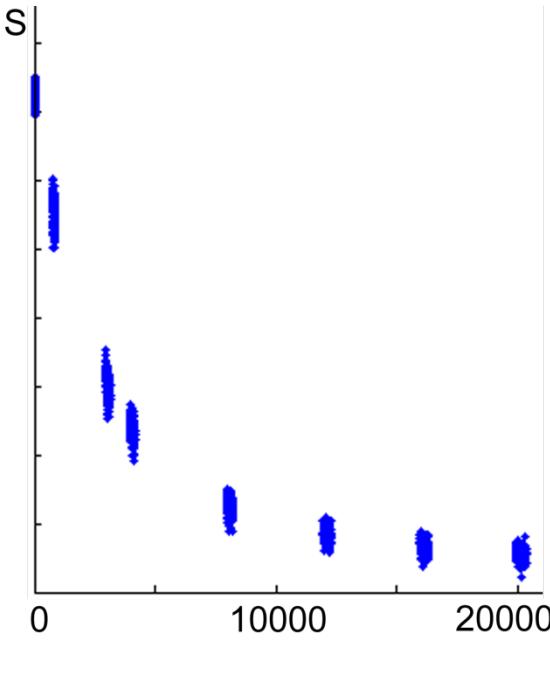


## Data impression

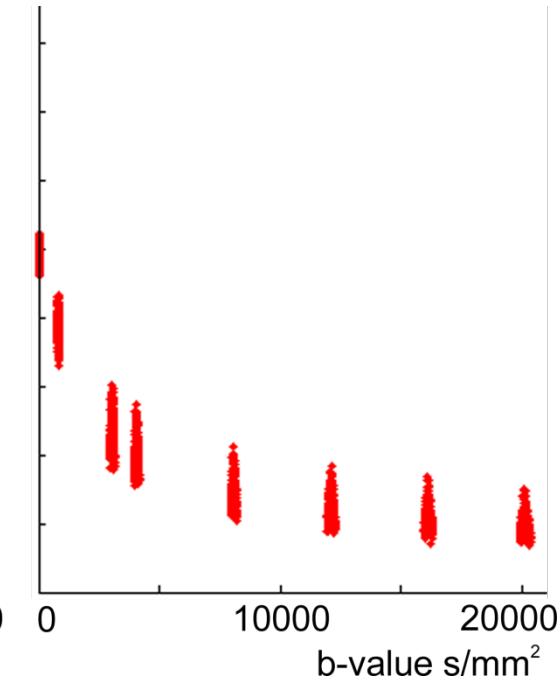
Normalized shell means



GM voxel

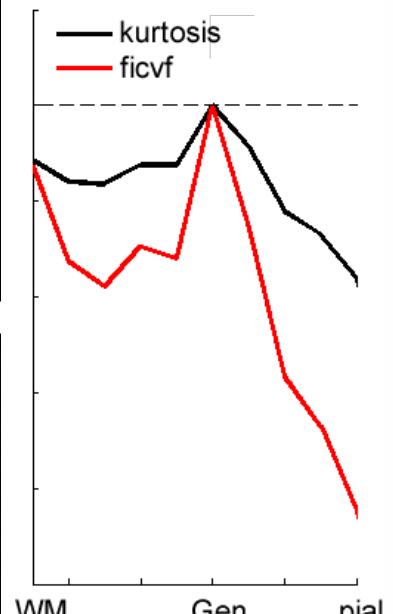
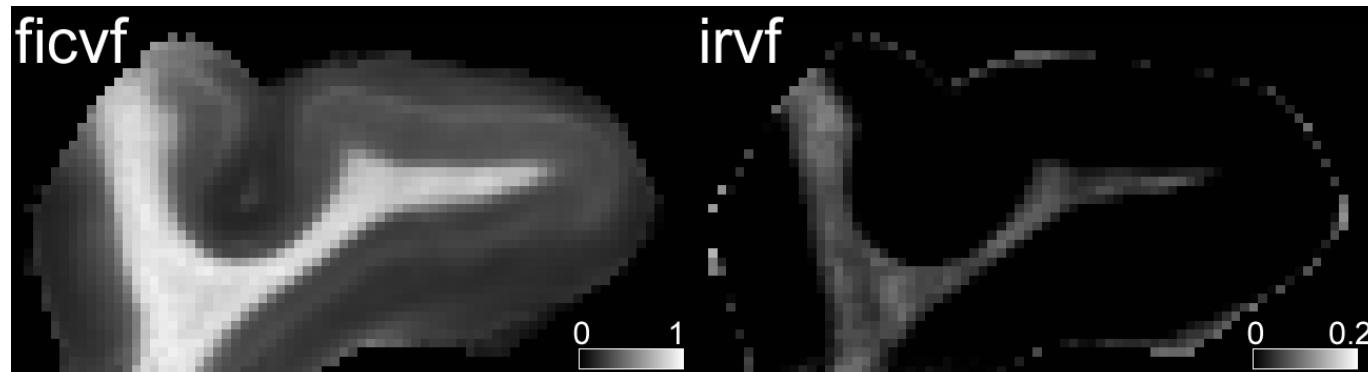


WM voxel



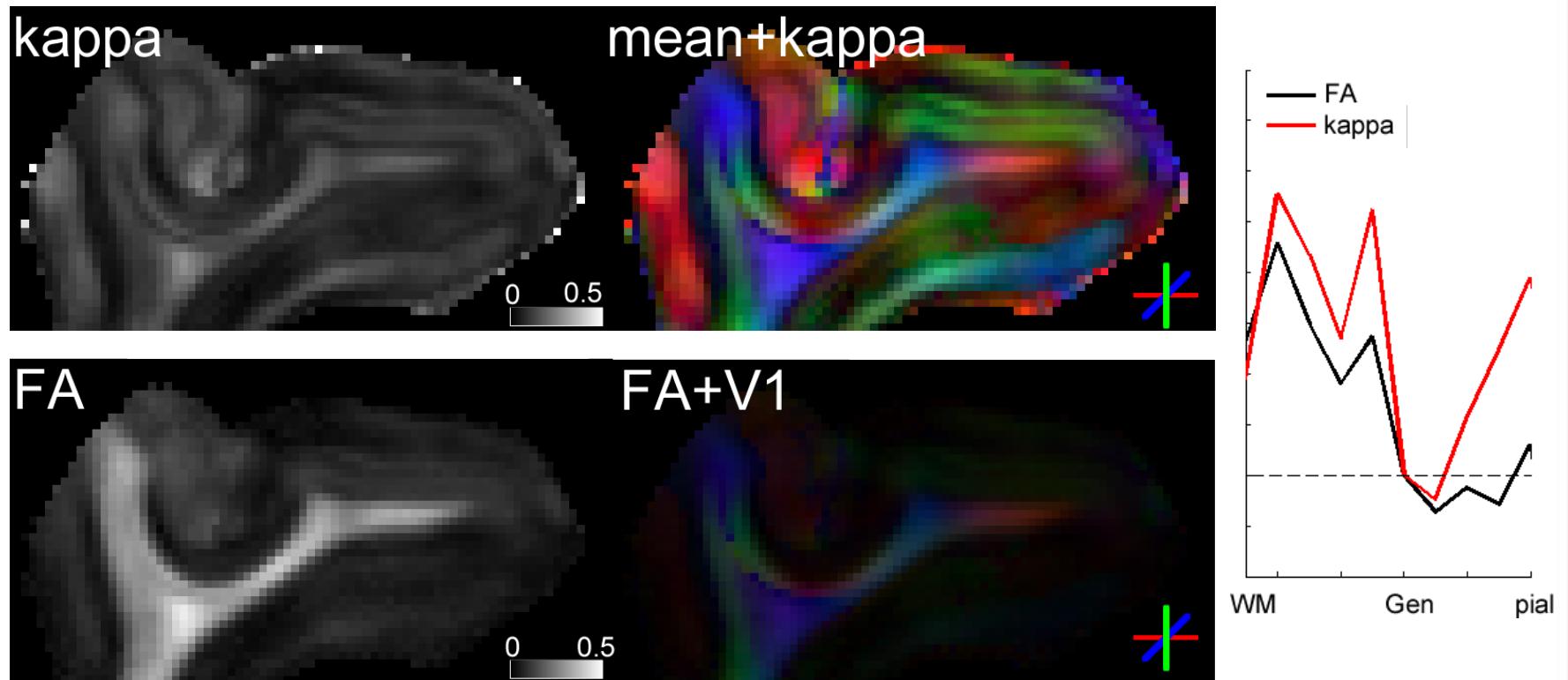


## Volume fractions



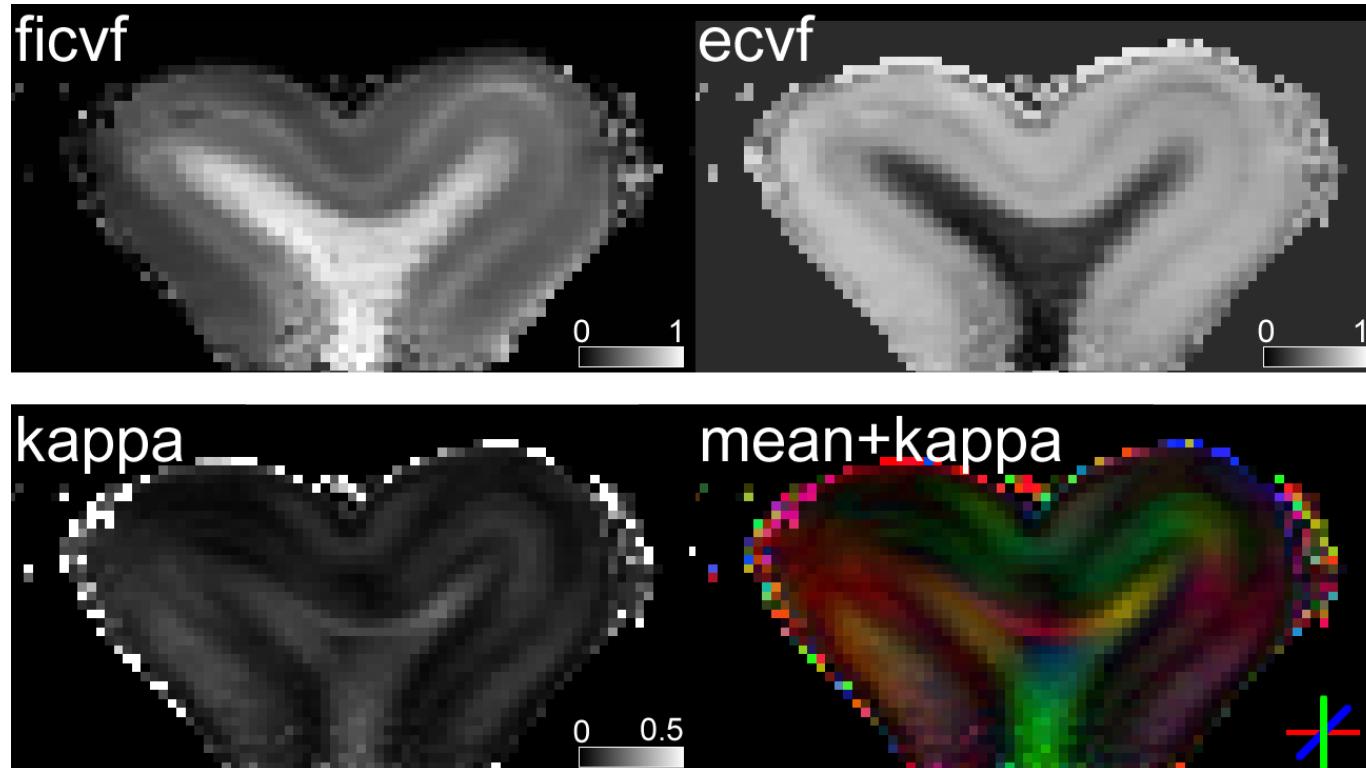


## Orientation dispersion



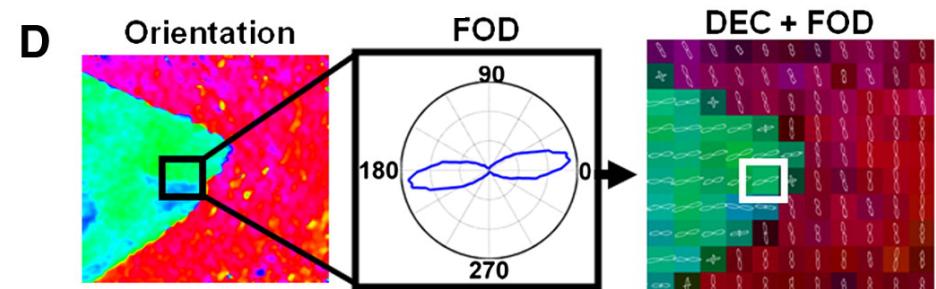
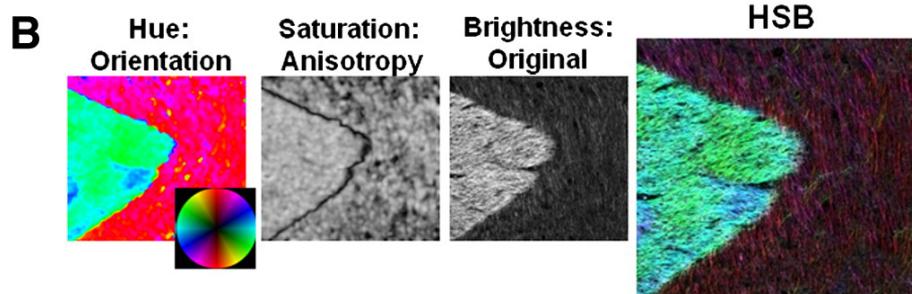
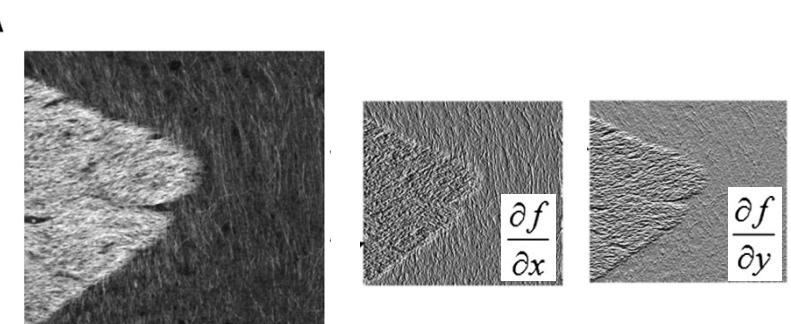


## Sample B (54 directions)



## Histology

- Samples bisected and embedded in paraffin
- Stains on consecutive 5 µm sections:
  - Hematoxylin (cell bodies) & Eosin (cytoplasm)
  - Luxol Fast Blue (myelin)
  - Bodian (axons)
- Virtual slice microscopy (20 X)
- Structure tensor analysis  
(Budde and Frank, 2012)



Tile 1

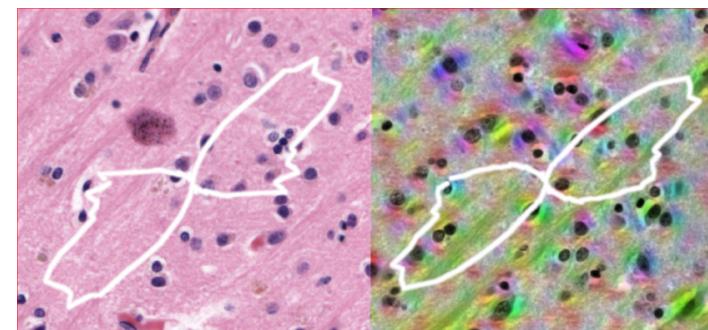
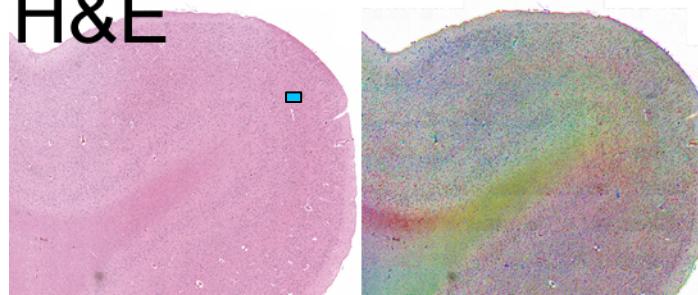
Tile 2



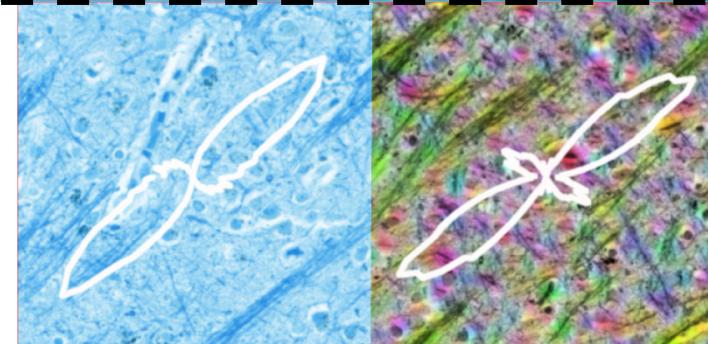
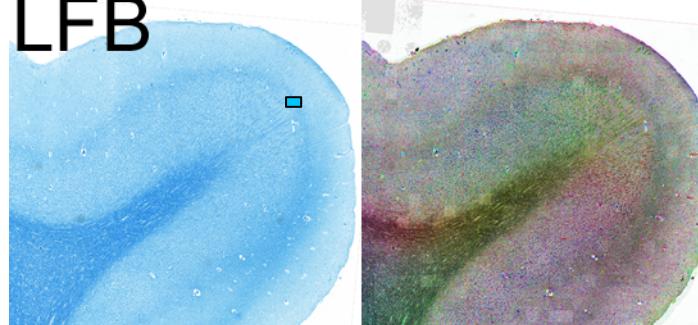
Histology



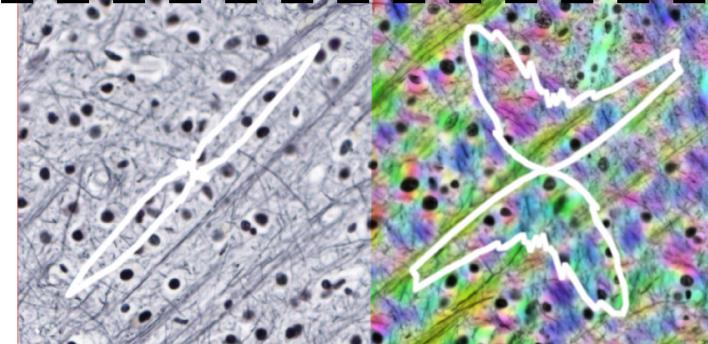
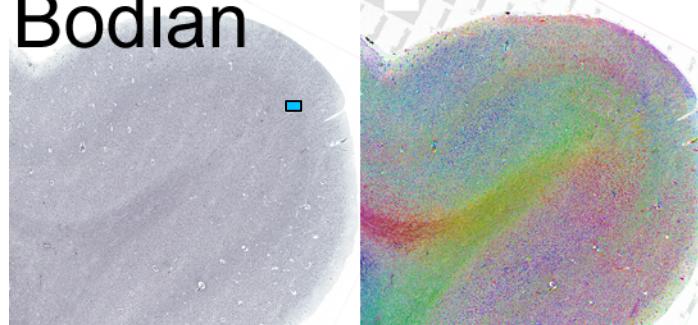
H&E



LFB



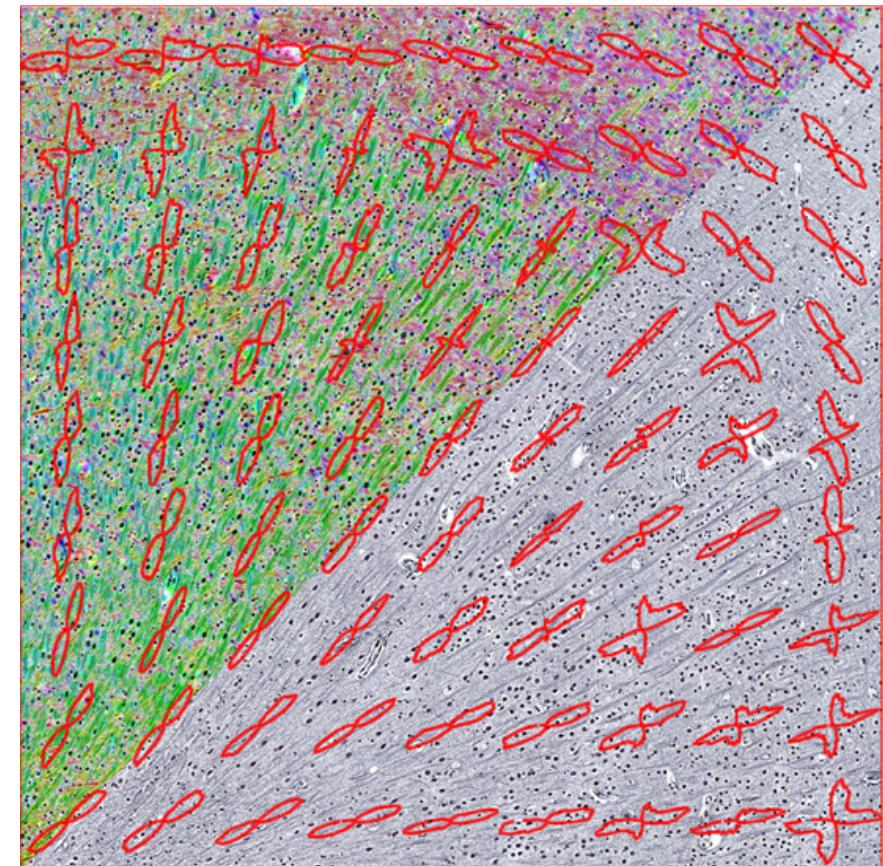
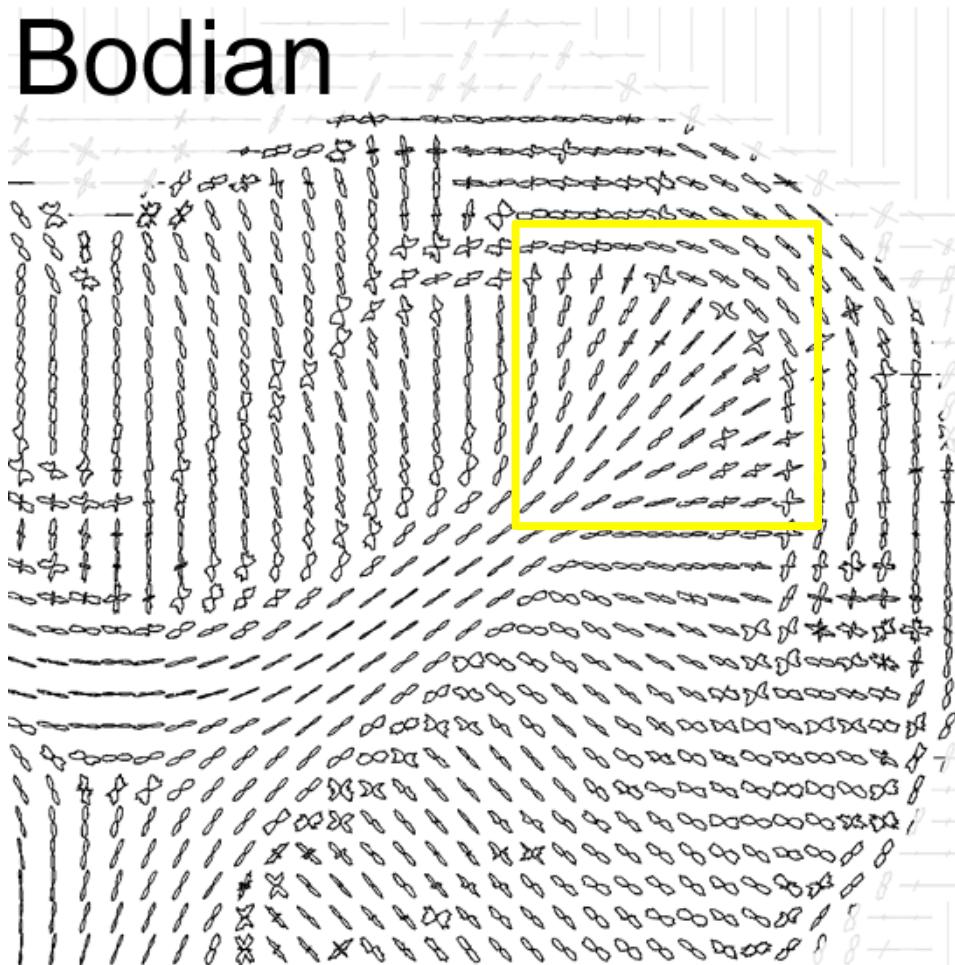
Bodian





## Histology

# Bodian





## Conclusions

### NODDI

- High layer discriminability
- Sharp delineation of layer boundaries in GM and WM
- Interpretable measures
- Cortical *in vivo* investigations feasible in clinical scan times

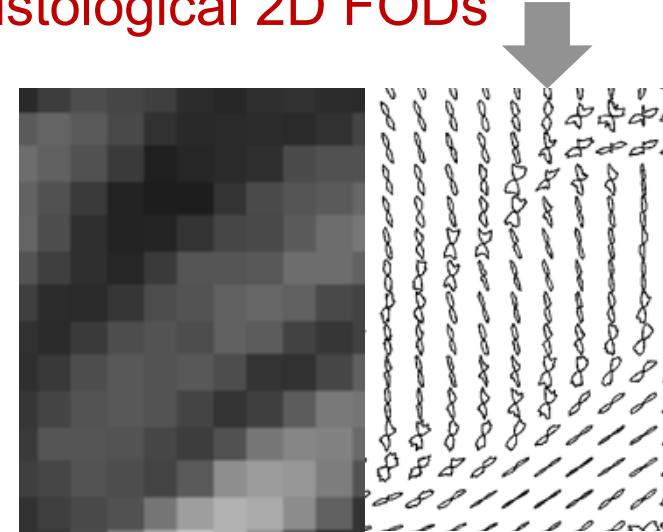
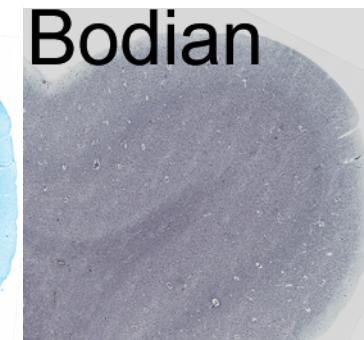
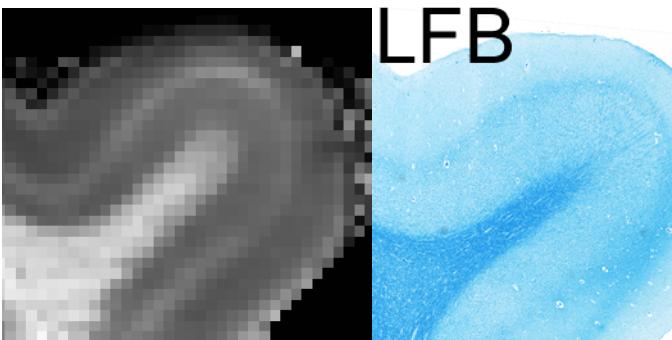


## Conclusions

### Histology

- Structure tensor useful in human, cortical histology

- Neurite volume fraction resembles myelin and axon stains
- Orientation dispersion reflected in histological 2D FODs



- no histological label for dendritic space included



# Thank you

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Pathology: Ine Mamor-Cornelissen

MPI Köln: Matthias Hoehn

DCCN: Markus Barth

Marcel Zwiers

**VIP Brain Networks**

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