# Validation of NODDI estimation of dispersion anisotropy in V1 of the human neocortex

Submission Number:

6482

# Submission Type:

Abstract Submission

#### Authors:

Maira Tariq<sup>1</sup>, Michiel Kleinnijenhuis<sup>2</sup>, Anne-Marie van Cappellen van Walsum<sup>3</sup>, Gary Zhang<sup>1</sup>

## Institutions:

<sup>1</sup>University College London, London, United Kingdom, <sup>2</sup>University of Oxford, Oxford, United Kingdom, <sup>3</sup>Radboud University Medical Centre, Nijmegen, Netherlands

## Introduction:

This work validates the estimation of neurite dispersion anisotropy in the brain, using Bingham-NODDI [1], an extension of the diffusion MRI technique called NODDI [2]. The original NODDI provides indices of neurite (axons and dendrites) morphology that are sensitive and specific to microstructural changes [3-7]. Bingham-NODDI additionally allows the estimation of neurite dispersion anisotropy, which can enhance the accuracy of tractography algorithms [8]. The in vivo feasibility of Bingham-NODDI has been evaluated in [1]. The present study validates its indices using high-resolution ex vivo imaging data of the human primary visual cortex (V1), a well characterised region of the neocortex known to include fibres that fan or bend into the cortical layers.

# Methods:

DATA: Diffusion weighted images were acquired for a fixed sample of the human V1 (as used in [9]), using a 9.4T small animal scanner. The acquisition included b=[0,1000,4000,8000,12000,16000,20000]s/mm<sup>2</sup> with  $\delta$ =8.4ms,  $\Delta$ =12.8ms, TE=27ms, and 60 gradient directions. The images were of 0.2mm isotropic resolution with FOV=28.8x28.8mm, covering the cortex and the underlying white matter (WM).

MODEL: NODDI is a multi-compartment model that accounts for the dispersion of neurites with an orientation density function (ODF). The original Watson-NODDI uses the Watson distribution [10] which can only model isotropic dispersion about the dominant orientation ( $\mu_1$ ); the Bingham-NODDI instead uses the Bingham distribution [10] and enables the modelling of anisotropic dispersion. Both models estimate the neurite density ( $v_{in}$ ),  $\mu_1$  and the orientation coherence ( $\kappa$ ) about  $\mu_1$ . Bingham-NODDI additionally estimates the primary dispersion orientation ( $\mu_2$ ) and the associated coherence parameter ( $\beta$ ). The eigenvalues of the scatter matrix of an ODF,  $\tau_1$  and  $\tau_2$ , represent the relative neurite concentrations along  $\mu_1$  and  $\mu_2$ , and are functions of  $\kappa$  and  $\beta$ . The dispersion anisotropy is quantified as: DAI=( $\tau_2$ - $\tau_3$ )/ $\tau_1$ , as described in [1].

FITTING: Both models were fitted to the data using the NODDI Matlab Toolbox. Watson-NODDI parameters were obtained using the optimisation procedure described in [2]; the intrinsic diffusivity was also fitted. The maximum-likelihood parameters from the Watson-NODDI fit were used to initialise the fitting for Bingham-NODDI. The parameters from the two models were evaluated using the Bayesian Information Criterion (BIC), a standard model comparison metric.

### **Results:**

PARAMETER EVALUATION: Fig.1 shows the maps of the Bingham-NODDI estimates,  $\beta$  and DAI. The DAI map shows highest values in WM that gradually get lower as the WM fibres disperse into the cortical grey matter (GM), signifying the presence of fanning and bending fibres in the cortical areas. The  $\beta$  plot has a contrast very similar to DAI, but the higher values are limited to WM regions, unlike the higher DAI values, showing that the WM fibres can be traced extending into the cortex using the DAI estimates.

MODEL COMPARISON: The BIC maps in Fig.2 demonstrate that Bingham-NODDI explains the data better than Watson-NODDI. Watson-NODDI fits the data particularly poorly in WM regions, especially near the WM/GM boundary, areas where dispersion anisotropy is expected. Fig.2 also shows that the  $\kappa$  maps from Bingham-NODDI have higher intensity in WM, as well as the various cortical layers, compared to Watson-NODDI. This increased intensity and contrast is more closely aligned with histology, as it is found in [9] that Watson-NODDI underestimates the actual neurite coherence.

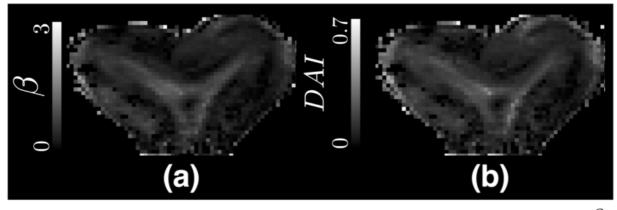
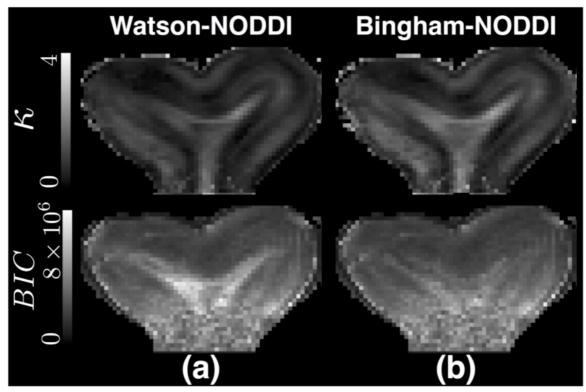


Fig.1 Maps of the novel parameters of Bingham-NODDI, (a)  $\beta$  and (b) DAI, for a slice of the primary visual cortex sample.



# Fig.2 Maps of $\mathcal{K}$ estimates and quality of fit (BIC) for the two NODDI models, for a slice of the primary visual cortex sample.

#### **Conclusions:**

We show that Bingham-NODDI is able to capture the cortical fibres known to exhibit fanning/bending [9], with a measure of dispersion anisotropy. This has implications for enhancing our understanding of the brain microstructure and connectivity, even in regions of complex cytoarchitecture, like the cortex.

## **Imaging Methods:**

Diffusion MRI<sup>2</sup>

#### **Modeling and Analysis Methods:**

Diffusion MRI Modeling and Analysis <sup>1</sup>

Keywords: MRI

NORMAL HUMAN

Other - NODDI, Diffusion MRI, Dispersion anisotropy, Bingham Distribution, Ex vivo

# $^{1\left| 2\right| }$ Indicates the priority used for review

Would you accept an oral presentation if your abstract is selected for an oral session?

Yes

Please indicate below if your study was a "resting state" or "task-activation" study.

## Other

Healthy subjects only or patients (note that patient studies may also involve healthy subjects):

## Healthy subjects

Internal Review Board (IRB) or Animal Use and Care Committee (AUCC) Approval. Please indicate approval below. Please note: Failure to have IRB or AUCC approval, if applicable will lead to automatic rejection of abstract.

## Not applicable

Please indicate which methods were used in your research:

Diffusion MRI

For human MRI, what field strength scanner do you use?

If Other, please list - 9.4T

### Which processing packages did you use for your study?

Other, Please list - NODDI toolbox, ITK-SNAP

FSL

#### Provide references in author date format

[1] Tariq, M. (2014), 'In vivo estimation of dispersion anisotropy of neurites using diffusion MRI', Proceedings of the Annual Meeting of the Medical Image Computing and Computer Assisted Intervention (MICCAI), Boston, USA.

[2] Zhang, H. (2012), 'NODDI: Practical in vivo neurite orientation dispersion and density imaging of the human brain', NeuroImage, Vol.61, pp.1000-1016.

[3] Winston, G. (2014), 'Advanced diffusion imaging sequences could aid assessing patients with focal cortical dysplasia and epilepsy', Epilepsy Research, Vol.108(2), pp.336-339.

[4] Kunz, N. (2014), 'Assessing white matter microstructure of the newborn with multi-shell diffusion MRI and biophysical compartment models', NeuroImage, Vol.96, pp.288-299.

[5] Owen, J.P. (2014), 'Aberrant White Matter Microstructure in Children with 16p11.2 Deletions', Journal of Neuroscience, Vol.34(18), pp.6214-23.

[6] Timmers, I. (2014), 'White matter microstructure pathology in classic galactosemia revealed by neurite orientation dispersion and density imaging', Journal of Inherited Metabolic Disease, pp.1-10.

[7] Lemkaddem, A. (2014), 'Connectivity and tissue microstructural alterations in right and left temporal lobe epilepsy revealed by diffusion spectrum imaging ', NeuroImage: Clinical, Vol.5, pp.349-358.

[8] Rowe, M. (2013), 'Beyond Crossing Fibers: Tractography Exploiting Sub-voxel Fibre Dispersion and Neighbourhood Structure', Information Processing in Medical Imaging, Lecture Notes in Computer Science, Vol.7917, pp.402-413.

[9] Kleinnijenhuis, M. (2014). 'Fibres in the gyrus: characteristics in ex vivo diffusion weighted imaging and histology', Imaging fibres in the brain. Ph.D. Thesis, Radboud University, Nijmegen, The Netherlands.

[10] Mardia, K.V. (1990), 'Directional statistics', Wiley series in probability and statistics, John Wiley & Sons, Ltd.