



# Extracting a biomarker for the mean cross-sectional area from the ODF

Rutger Fick, Gonzalo Sanguinetti, Rachid Deriche

## ► To cite this version:

Rutger Fick, Gonzalo Sanguinetti, Rachid Deriche. Extracting a biomarker for the mean cross-sectional area from the ODF. Human Brain Mapping, Jul 2014, Hamburg, Germany. <hal-01095197>

**HAL Id: hal-01095197**

**<https://hal.inria.fr/hal-01095197>**

Submitted on 15 Dec 2014

**HAL** is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.

## Extracting a biomarker for the mean cross-sectional area from the ODF

---

**Submission Number:**

688

**Authors:**Rutger Fick<sup>1</sup>, Rachid Deriche<sup>1</sup>, Gonzalo Sanguinetti<sup>2</sup>**Institutions:**<sup>1</sup>INRIA, Sophia-Antipolis, France, <sup>2</sup>INRIA, Sophia Antipolis, France**First Author:**

*Rutger Fick* - Lecture Information | Contact Me  
INRIA  
Sophia-Antipolis, France

**Introduction:**

Finding new biomarkers related to the microstructure of white matter (WM) is an active area of research in the MRI community [1]. As opposed to the usual MRI markers such as fractional anisotropy (FA), these biomarkers provide a closer insight on the tissue structure. We introduce a new microstructure based biomarker that is related to the axon diameter distribution (ADD) and can be obtained with a q-space imaging [2] technique like DSI [3] or MAP [4]. This feature is related with the nature and purpose of WM paths in both normal and pathological conditions [5], and is obtained from the Orientation Distribution Function (ODF) [6] as twice its maximum value. We show that this value is related with the mean cross-sectional area (MCSA) of an ensemble of parallel axons. The same geometric feature was proposed in [4] as a scalar index of microstructure, but was not related to the ODF. In this work we give the formal relation between this microstructure feature and the ODF, and validate it using state-of-the-art numerical simulations.

**Methods:**

In anisotropic WM voxels where the underlying tissue is made of aligned axons, the angle of the ODF maxima coincides with the axon orientation. In [4] it was proved, under the hypothesis that the gradient pulses ( $\delta$ ) are infinitely small and we are under the long diffusion time regime ( $\Delta \gg \delta$ ), that the integral of the q-space attenuation over the plane perpendicular to a fiber path is equal to the reciprocal of the MCSA of the ADD. Using the central section theorem of the 3D Fourier transform, in [7] it was proved that this integral is equivalent to twice the amplitude of the ODF in the direction normal to the plane. Combining both results, we obtain the relation between the MCSA and the ODF amplitude.

To validate this relationship, we generate synthetic diffusion data using Monte-Carlo simulations with Camino [8]. In the simulations, WM is a 3D environment called substrate, formed by randomly placed parallel cylinders. A total of 22 different substrates were considered, each one containing 10,000 cylinders whose diameter are randomly taken from a Gamma distribution that approximates the ADDs observed in histology data [9]. A DSI acquisition protocol was used to sample 515 q-space points in a cubic fashion with q-values ranging between 0 and 432 1/mm with  $\delta=1\text{ms}$ ,  $\Delta=100\text{ms}$  and  $TE=.101\text{ms}$ . To ensure convergence, we set a large number of particles (3,000,000). To be consistent with our model, we only consider the signal produced by the intra-axonal space. Some of the substrates simulated with Camino and their corresponding ADDs are visualized in Figures 1 and 2. In the histograms, the solid line represents the analytic Gamma distribution as a function of axon radius ( $r$ ), while the bars represent the simulated radii distribution.

**Results:**

Figure 3 shows the known versus the estimated MCSA in  $\mu\text{m}^2$  (denoted as  $\hat{M}$ ) for 22 substrates. The MCSA was estimated as twice the amplitude of the ODF, calculated as in [6]. The estimated values correlate well with the known values, but are slightly overestimated. This can be attributed to the truncation of the q-space by the sampling scheme, causing an underestimation of the q-space plane integral, and opposite for the MCSA.

**Conclusions:**

We introduced an easy to calculate biomarker that can characterize the axon diameter distribution and can be obtained from any QSI acquisition as the ODF amplitude. We validate the assumptions by means of Monte-Carlo simulations of WM-like environments. Validations on real data are to come.

**Modeling and Analysis Methods:**

Diffusion MRI Modeling and Analysis

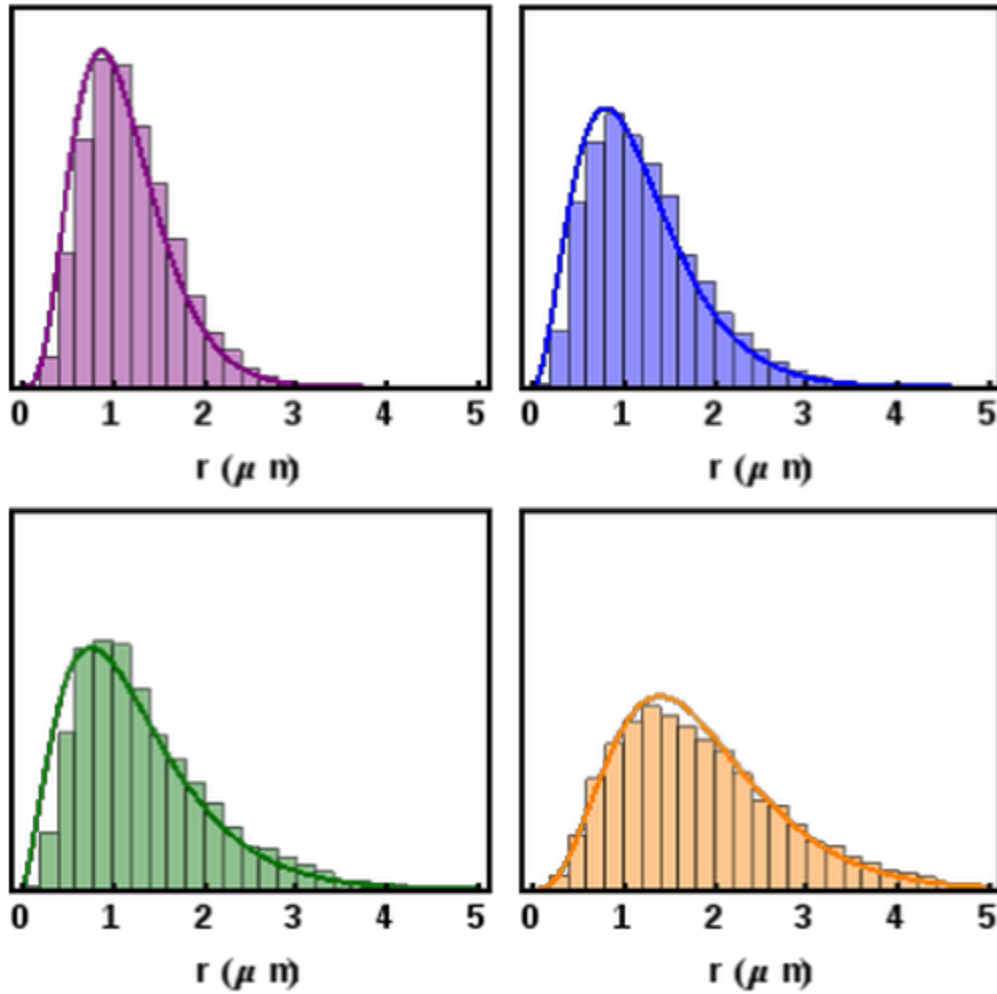


Figure 2. Analytical and simulated Gamma distributions as a function of radius.

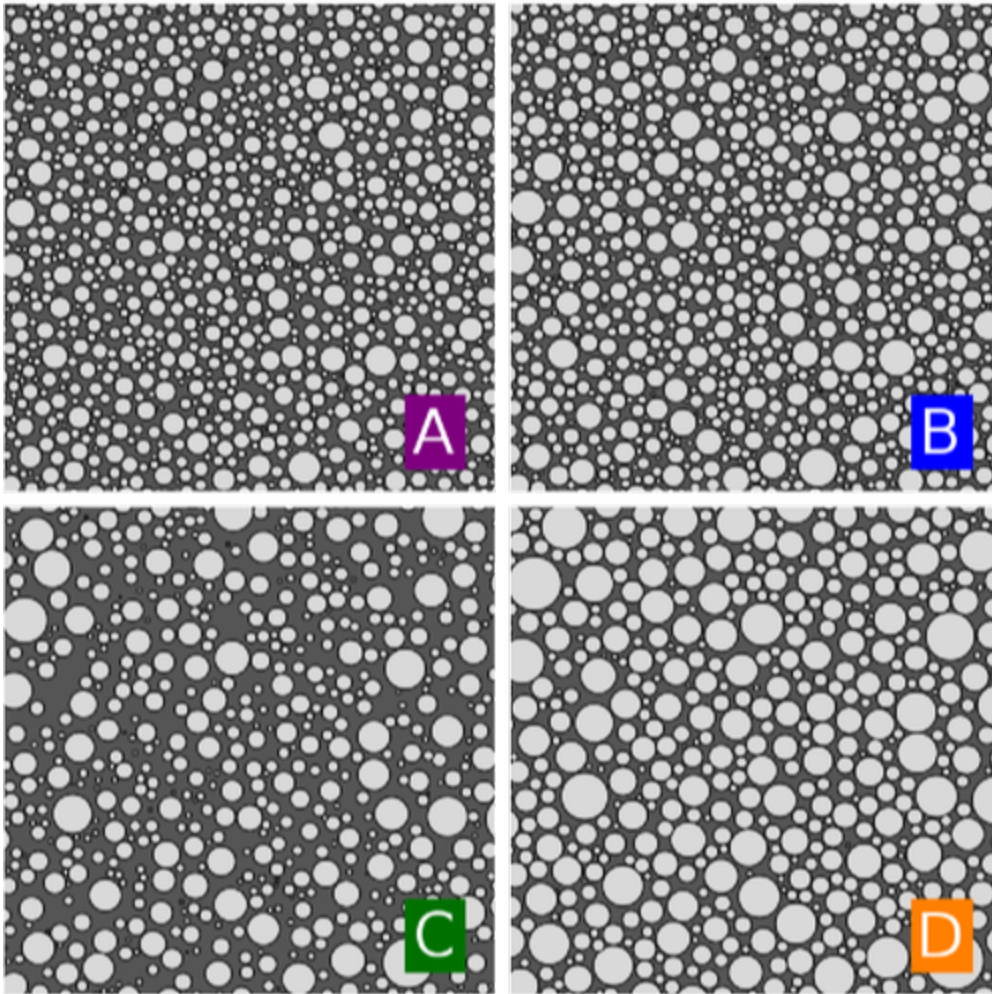


Figure 1. Cross-section of simulated radii distributions

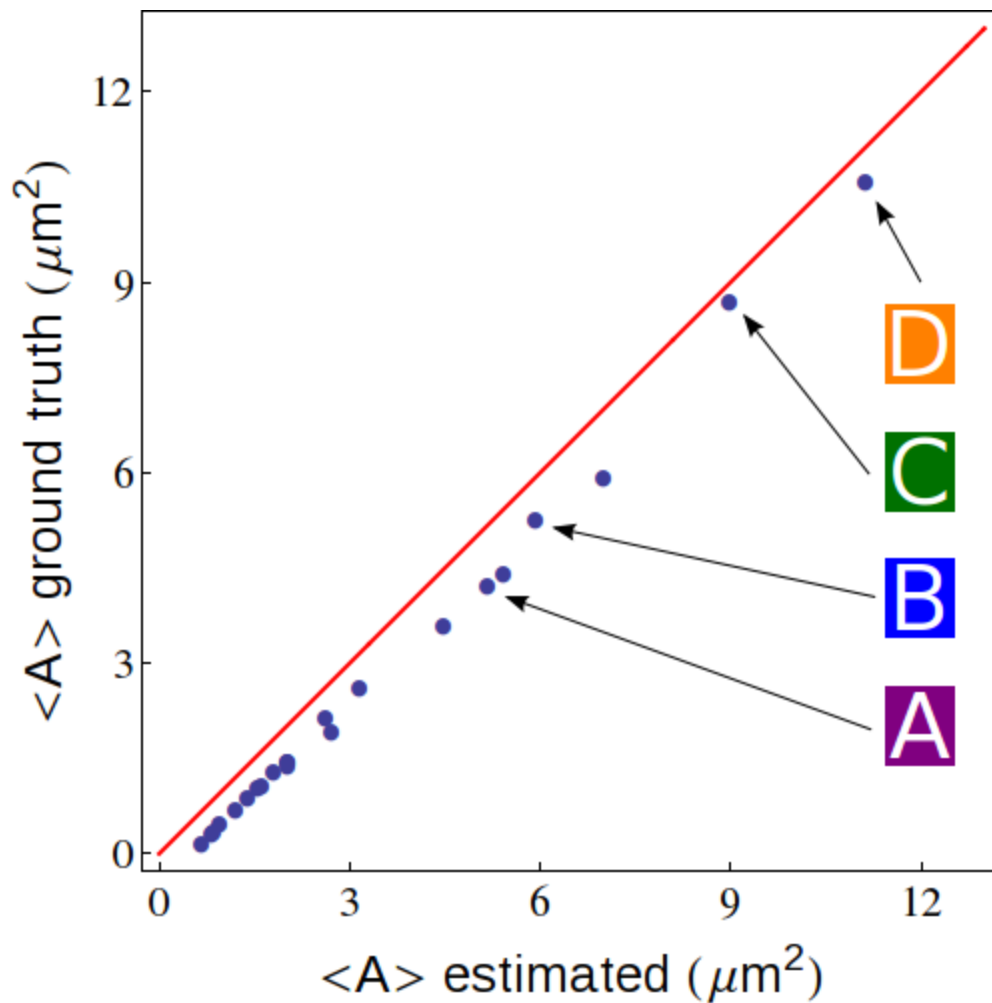


Figure 3. Comparison of known versus estimated mean cross-sectional area for 22 distributions.

#### Reference

- [1] Assaf, Y (2013), 'The CONNECT project: Combining Macro-and Micro-structure', *NeuroImage*, vol. 80, pp. 273-282.
- [2] Callaghan, P. (1990), 'High-resolution q-space imaging in porous structures', *Journal of Magnetic Resonance*, vol. 90, pp.177-182.
- [3] Wedeen, V. (2005), 'Mapping complex tissue architecture with diffusion spectrum magnetic resonance imaging', *Magnetic Resonance in Medicine*, vol. 54.6, pp. 1377-1386.
- [4] Ozarslan, E. (2013), 'Mean apparent propagator (map) MRI: A novel diffusion imaging method for mapping tissue microstructure', *NeuroImage*, vol. 78, pp. 16-32.
- [5] Hughes, J. (2007), 'Autism: the first firm finding = underconnectivity?', *Epilepsy & Behavior*, vol. 11.1, pp. 20-24.
- [6] Tuch, D (2004), 'Q-ball imaging", *Magnetic Resonance in Medicine*, vol. 52.6, pp. 1358-1372.
- [7] Tristan-Vega, A. (2010), 'A new methodology for the estimation of fiber populations in the white matter of the brain with the Funk-Radon transform', *NeuroImage*, vol. 49.2, pp. 1301-1315.
- [8] Alexander, D. (2008), 'A general framework for experiment design in diffusion MRI and its application in measuring direct tissue-microstructure features', *Magnetic Resonance in Medicine*, Vol. 60, pp. 439-448.
- [9] Alexander, D. (2010), 'Orientationally invariant indices of axon diameter and density from diffusion MRI', *NeuroImage*, vol. 52.4, pp. 1374-1389.