Measuring accurate small compartmental dimensions using double-PFG NMR: from water-filled microcapillaries to biological cells

Noam Shemesh¹, Evren Özarslan², Peter J Basser² and Yoram Cohen¹

¹School of Chemistry, The Raymond and Beverly Sackler Faculty of Exact Sciences, Tel Aviv University, Israel.

²Section on Tissue Biophysics and Biomimetics, NICHD, National Institutes of Health, Bethesda, Maryland, USA

Obtaining microstructural information such as pore size, anisotropy, and shape from opaque specimens is important in a variety of disciplines ranging from material science to neuroscience. Diffusion-NMR and especially the single-pulsed-field-gradient (s-PFG) methodologies (Fig. 1A) are useful for characterizing anisotropy in coherently-packed anisotropic pores, and the compartment size can be extracted from diffusion-diffraction minima if the pores are monodisperse. When the specimen is characterized by size distributions, the q-space approach can be used to infer on relative compartment sizes. Howeover, the s-PFG approaches require extremely strong gradient amplitudes to extract compartment dimensions, and are very limited when orientation distributions are present.

The double-PFG (d-PFG) methodology (Fig. 1B) is an extension of s-PFG that employs a second PFG pair. The angular-d-PFG experiment, in which the angle between the gradient pairs is varied, has gained interest owing to recent theoretical studies prediting that angular-d-PFG can afford new microstructural information even at low gradient amplitudes, and even when the pores are randomly oriented or have broad size distributions.

Therefore, we conducted angular-d-PFG experiments on water-filled microcapillaries with inner-diameter (ID) of $5\pm1~\mu m$. Figure 2 shows the experimental points (symbols) and theoretical fits (lines) of the angular-d-PFG experiment. Importantly, a very accurate compartment size can be extracted from the bell-shaped functions using weak gradients (as low as 15 G/cm). For reference, the s-PFG could only extract such size using gradient strengths of 573~G/cm.

Recently, these findings were applied to randomly oriented yeast cells. The cell size extracted from d-PFG measurements was $5.46\pm0.46~\mu m$, whereas light microscopy measurements yielded $5.32\pm0.32~\mu m$. Such findings may become important in clinical d-PFG MRI, as well as in accurately characterizing porous media.

