

## **Advanced Diffusion Physics Applied to Body Imaging**

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Diffusion-weighted imaging (DWI) offers insight into cellular status, density, and structural organization by way of sequences sensitive to water mobility which is affected by these properties [1, 2]. Macroscopic tissue motion unrelated to diffusion can confound in vivo diffusion measurements therefore single-shot techniques are nearly exclusively used to mitigate extraneous motion, particularly in the abdomen. Microcirculation flow through randomly-oriented capillaries in the presence of diffusion-sensitization gradients will also appear as a hyper diffusion-like attenuation in the low b-value regime [3, 4]. In the high b-value extreme where diffusion-based contrast is high, quantitative measurement of signal attenuation with increasing b-value is susceptible to noise limitations [5]. While not as pronounced as in neuro tissue, water mobility in body tissues may be directional due to true underlying cyto-architecture or appear anisotropic due to residual bulk motion artefact. Unlike DWI of the brain, an effective fat suppression method is crucial for successful body DWI sequences. In summary, diffusion imaging of the body requires organ site-specific customization of protocols to deal with: perfusion contamination at low b-values, SNR limitations in the high b-value regime, shim quality and fat suppression over a large FOV, multi-axis measurements to properly quantify isotropic diffusion (eg. ADC) or anisotropic features, as well as possible synchronization with cardiac/respiration to reduce residual bulk motion errors. Recent technical acquisition enhancements have successfully addressed many of these issues such that DWI of the body has gained rapid growth in a variety of applications.

Proper selection of b-values and gradient directions depend on the given body DWI application and objective [6]. Diffusion sensitization pulses on at least three orthogonal gradient axes are required to quantify a rotationally-invariant diffusion coefficient (ie, ADC or mean diffusion) [7, 8]. If anisotropic diffusion indices are sought (such as fractional anisotropy), at least 6 non-collinear directions are required although 9-16 directions are not uncommon. The quantity and range of acquired b-values also should be suited for the given application and signal quality properties. If simple quantification of ADC is desired, only two b-values are required as is typical in most clinical DWI studies to date. However, if one seeks to disentangle potential perfusion effects from molecular diffusion, or to detail true biophysical multi-exponential diffusion features then additional b-value samples are needed to fit DWI signals to a specified multi-exponential model. For example, diffusion may be modelled as a bi-exponential decay where characterization of “fast” and “slow” diffusion components, and their fractional contribution requires additional b-values to fit at least three model coefficients [9, 10]. Alternatively, the stretched-exponential models a continuum of diffusion decays embodied in one “distributed diffusion coefficient” and involves only two model coefficients [11]. Perfusion influences are particularly relevant to diffusion measurements in vascular-rich lesions/tissues. In such instances, signals acquired in the low b-value range (eg. 0 to 150s/mm<sup>2</sup> ) are affected by perfusion. It is empirically challenging to extract the “perfusion fraction” from measurements over the low and high b-value regime, although these concepts are being revisited [4]. Alternatively, one may effectively extinguish perfusion signals and their influence by only including b-values above 150s/mm<sup>2</sup> in diffusion calculations. The maximum b-value should be set such that signals recorded at that b-value are adequately above the noise floor. This maximum b-value depends on the SNR achievable for the target organ/tissue and the water diffusion coefficient of these tissues – the lower the diffusion value, the higher the b-value achieved before the signal approaches the noise floor. Field strength, receiver coil, acquired resolution and scan time provide some operator control to improve SNR and DWI quality, although reasonable guidelines for several body DWI protocols have been suggested [6].

One body DWI application that may have practical clinical value is lesion detection in a limited anatomical region, or over a “whole-body” survey scan [12, 13]. Pathology characterized by relatively long T2 appear hyper-intense on DWI due, in part, to inherent high T2-weighting in DWI. Other normal long-T2 tissues such as blood and fluids, would also appear bright if not for incremental diffusion sensitization that attenuate high mobility fluid signals. Therefore lesions having long T2 and moderate to low water mobility, such as is common in solid lesions tend to exhibit conspicuous hyper intensity on moderate to high b-value DWI. Interestingly, despite their conspicuous hyper intense DWI signal, water mobility via ADC of these lesions often is not lower than normal tissues. If only “lesion detection” is desired, collection of multiple b-values is not strictly required. However, other applications such as lesion/tissue characterization or treatment response assessment via ADC, or multi-exponential analysis of DWI decays, or anisotropy study of organized tissues, then additional DWI conditions are required with commensurate increases in scantime. In this lecture, principles of body DWI measurements, protocol design, and biophysical models for analysis will be presented.

### References:

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