# Time dependence of the diffusional kurtosis in the human calf muscle

Anja Maria Marschar<sup>1</sup>, Tristan Anselm Kuder<sup>1</sup>, Bram Stieltjes<sup>2</sup>, and Frederik Bernd Laun<sup>1,2</sup>

<sup>1</sup>Department of Medical Physics in Radiology, German Cancer Research Center (DKFZ), Heidelberg, Germany, <sup>2</sup>Quantitative Imaging Based Disease Characterization, German Cancer Research Center (DKFZ), Heidelberg, Germany

# Introduction

The diffusional kurtosis ( $K_{app}$ ) describes the deviation of the diffusion propagator in tissue from a Gaussian function [1]. It is usually measured in combination with the apparent diffusion coefficient ( $D_{app}$ ) and yields additional information on the tissue structure in the human body. It has been measured in several organs, for example in the brain [2] and prostate [3], and was shown to be a valuable parameter to describe diseased tissue, e.g. in prostate cancer [3]. The aim of this work was to perform measurements of  $K_{app}$ , including its dependence on the diffusion time, in healthy muscle tissue to establish a basis of comparison for the investigation of muscle diseases, e.g., muscle dystrophy.

## **Materials and Methods**

A stimulated echo acquisition mode sequence [4] was used to measure diffusion weighted images (DWIs) in the human calf muscle at 1.5 Tesla (Magnetom Avanto, Siemens Medical Solutions, Erlangen, Germany). The parameters were TE = 63 ms (for mixing time TM = 100 ms) or TE = 46 ms (for TM = 500 ms) or TE = 38 ms (for TM = 1000 ms), TR = 2.8 s, BW = 1804 Hz/px, matrix 64 x 64, resolution 6.9 mm, slice thickness 10 mm, 3 averages. Three orthogonal diffusion gradient directions were used with the following b-values: 0, 500, 1000, 1500, 2000, 2500, 3000, 3500, 4000, 4500, 5000, 6000 s/mm<sup>2</sup> (for TM = 100 ms); 0, 1000, 2000, 3000, 4000, 5000, 5500, 6000, 6500, 7000, 7500, 8000, 8500, 9000, 9500, 10000 s/mm<sup>2</sup> (for TM = 500 and 1000 ms). D<sub>app</sub> and K<sub>app</sub> values were fitted with the following formula as proposed in [1] with the background noise  $\eta$ : S =  $\sqrt{\eta^2 + \left(S_0 \exp\left(-bD_{app} + \frac{1}{6}b^2D_{app}^2K_{app}\right)\right)^2}$ . The DWIs were smoothed with a

noise  $\eta$ :  $S = \sqrt{\eta^2 + (S_0 \exp(-bD_{app} + \frac{1}{6}b^2D_{app}^2 K_{app}))}$ . The DWIs were smoothed with a Gaussian filter with a half width of 1.5 voxels. For the quantitative evaluation, regions of interests were placed in the muscle tissue omitting regions near skin, bones and vessels.

### Results

Fig. 1 shows maps of  $D_{app}$  and  $K_{app}$ , which both depend on the diffusion direction. Fig. 2 and 3 show the time-dependence of the ROI-averaged values of  $D_{app}$  and  $K_{app}$ . The z-direction is approximately parallel to the fiber direction. In z-direction the  $D_{app}$ -values are higher and the  $K_{app}$ -values are smaller compared to x- and y-direction.  $D_{app}$  decreases moderately with increasing TM while  $K_{app}$  decreases strongly.

### Discussion

The drop of  $D_{app}$  with increasing diffusion time was already observed in [5] and absolute values are in agreement with the values presented in [5]. To our knowledge, the time dependence of the kurtosis in muscle tissue has not been investigated to date. Interestingly, a much larger anisotropy can be observed for  $D_{app}$  than for  $K_{app}$ .  $K_{app}$  is almost isotropic. For small mixing times (100 ms), the diffusion process is mainly located in either the intracellular or the extracellular compartment, as muscle cells typically have a diameter of about 50 µm [6]. At this time scale, the myofibrils of about 1 µm [6] diameter are not detectable, but the cell membrane causes a restriction. At longer mixing time (1000 ms), the kurtosis decreases, which may be attributed to an effectively free water exchange between intra- and extracellular space at this long time scale. Compared to a kurtosis value of 1.1 in white matter tissue of the brain [2], the kurtosis in muscle tissue is much smaller. There seems to be a fundamental difference between brain tissue and muscle tissue. In brain tissue, for small diffusion time, it is probably justified to assume the presence of two well-separated compartments with different diffusion coefficients, caused by the hydrophobic myelin sheet, which results in large  $K_{app}$ -values [1]. In muscle tissue, however, such an almost impermeable layer is not present, and this may well explain the low  $K_{app}$  in muscle when compared to white matter.

#### References

[1] J.H. Jensen *et al.*, Magnet. Reson. Med. **53** (2005); [2] H. Lu *et al.*, NMR Biomed. **19** (2006); [3] A.B. Rosenkrantz *et al.*, Radiology, **264** (2012);
[4] E.L. Hahn, Phys. Rev. **80** (1950); [5] E. Fieremans *et al.*, Proceedings of the ISMRM 19<sup>th</sup> Annual Meeting, Montreal **19**, 1153 (2011); [6] B. Alberts *et al.*, Garland Publishing (1994)



**Fig.1:** Maps of  $D_{app}$  and  $K_{app}$  in the human calf muscle for different mixing times TM and diffusion directions. The z-direction is oriented along the body symmetry axis in the lab system.



**Fig.2:** Time dependence of  $D_{app}$  in the human calf muscle.



**Fig.3:** Time dependence of  $K_{app}$  in the human calf muscle.