

### 3D-maps of T1 and magnetization transfer (MT) related saturation from MT-FLASH images

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#### Introduction:

Quantitative magnetization transfer (MT) measurements in brain are commonly performed using 3D-FLASH (1) with interleaved saturation of bound spins and continuous wave models. Employing the binary spin bath (BSB) to model the saturation transfer after MT-pulse in brain tissue, we showed that the evolution of the BSB comprises slow relaxation of both pools and rapid MT preserving magnetization (2). Here, the MT-FLASH signal was described by the free-evolution BSB model. Pertinent approximations permit to calculate T1 and MT-related saturation from an MT and two reference FLASH images.

#### Theory:

For  $TR/T1 \ll 1$  and small flip angle  $\alpha \ll 1$ , the FLASH signal can be simplified by approximating the exponential and trigonometric terms:

$$S = S_0 \alpha TR/T1 / [TR/T1 + \alpha^2/2]. \quad [1]$$

Like in the trigonometric form, the Ernst angle is defined by equality of saturation created by excitation (by  $1 - \cos \alpha \approx \alpha^2/2$ ) and saturation restored by subsequent relaxation (by  $1 - \exp(-TR/T1) \approx TR/T1$ ). MT from the bound macromolecular pool (subscript 'm'; saturated by a fraction of  $\delta_m$ ) to the observed bulk water (subscript 'f'; saturated by  $\delta_f \ll \delta_m < 1$ ) may be expressed as time-dependent increase,  $\delta_{MT}$ , in the observed saturation (3):

$$\delta_{obs}(t) = \delta_f + \delta_{MT}(t) = \delta_f + F (\delta_m - \delta_f) [1 - \exp(-R_T t)] / [1 - (1 - \delta_m) \exp(-R_T t)], \quad [2]$$

where  $F = M_m^0 / (M_f^0 + M_m^0)$  is the bound pool fraction and  $R_T$  is the observed transfer rate. Saturation by MT- and excitation pulse and intervals of free evolution were concatenated using the matrix formalism of (2). Discarding products of small terms ( $TR/T1$ ,  $\alpha^2$ ,  $\delta_f$ ,  $F$ ) in the general solution eliminated the dependence on intervals of TR. Thus, the additional saturation originating from the MT-pulse,  $\delta_{obs}(t)$ , appears in the signal equation:

$$S = S_0 \alpha TR/T1 / [TR/T1 + \alpha^2/2 + \delta_{obs}(TR)] \quad [3]$$

The 3 parameters in Eqs. [1] and [3],  $S_0$ ,  $T1$ ,  $\delta_{obs}(TR)$ , may be calculated by arithmetic operations from 3 images (MT + 2 references of different  $\alpha$ ).

#### Methods:

3D-FLASH MRI with non-selective excitation was performed at 3 Tesla (Siemens Magnetom Trio; 8-channel receive headcoil) on healthy adult volunteers and one Multiple Sclerosis (MS) patient. Sequence parameters were varied to validate Eq. [3] and optimize the SNR of the parameter maps. PD-weighting ( $\alpha = 5^\circ$ ) ensuring sufficient CSF signal was combined with T1-w ( $\alpha = 15^\circ$ ;  $TR = 11$  ms). Optimal MT-effect within SAR limits was achieved on PD-w FLASH ( $\alpha = 5^\circ$ ;  $TR = 25$  ms) by a 12.8 ms Gaussian MT-pulse ( $530^\circ$ ; 2.2 kHz offset) (4). 1.25 mm isotropic resolution and an iPAT factor of 2 yielded a compromise between SNR and a measuring time of 13 minutes. FSL 3.2 ([www.fmrib.ox.ac.uk/fsl](http://www.fmrib.ox.ac.uk/fsl)) was used for co-registration, brain extraction, calculation and display of maps, and histogram analysis.

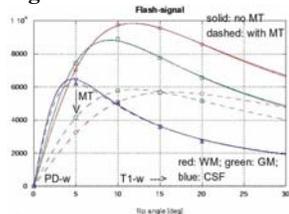
#### Results:

The fitted flip angle dependence with and without MT at  $TR = 25$  ms in Fig. 1 illustrates the signal drop due to additional MT-saturation compared to PD-FLASH in the caudate nucleus (GM) and splenium of the corpus callosum (WM), but not in lateral ventricle (CSF).  $\delta_{obs}$  increased with TR and pulse power. The native MT-w FLASH contrast of chronic MS black holes (right hemisphere) and an acute lesion with edema (left hemisphere) is shown in Fig. 2A. The reason for the T2-like appearance is that  $\delta_{obs}$  (about 1.5% in GM, 3% in WM; Fig. 2C) exceeded the 0.4% saturation due to  $\alpha$ . T1-values were slightly higher than reported for 3 T. Histogram analysis confirmed the high contrast visible on the  $\delta_{obs}$ -map (Fig. 2B) in comparison to the T1- and  $S_0$ -maps (Fig. 2D,E). Note the distinct difference between core and surrounding edema of the ring lesion, and the radial intensity gradient of the ring lesions.

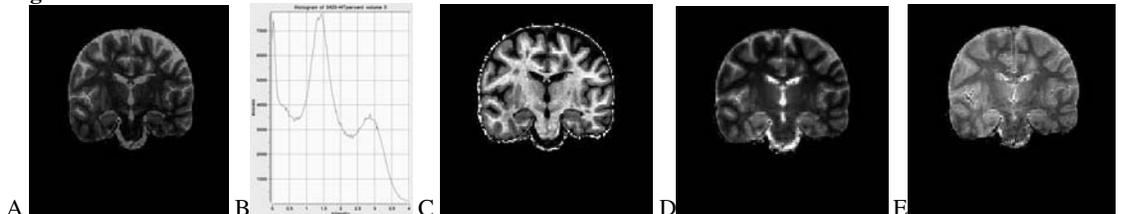
#### Discussion:

The calculated maps can provide surrogate parameters for tissue destruction (T1) and myelination ( $\delta_{obs}$ ) in MS.  $\delta_{obs}$  has a well defined physical meaning, namely the additional percentage saturation imposed by the MT-pulse. If direct effects can be neglected ( $\delta_{obs} = 0$ ),  $\delta_{obs}$  is proportional to the bound pool fraction, although still it depends on TR and MT-pulse. Its contrast properties and the possibility of obtaining high-resolution maps within clinically feasible time suggest it as an alternative to full MT-quantification and semi-quantitative MT-ratios. Further improvements in terms of SNR/measuring time/spatial resolution appear possible in combination with a 3D-multi-echo FLASH sequence.

#### Figure 1:



#### Figure 2:



Fitted flip-angle dependence MT-w FLASH contrast in MS patient (A), whole brain histogram of  $\delta_{obs}$  (B), and parameter maps of  $\delta_{obs}$ , T1,  $S_0$  (C-E)

#### References:

(1) Frahm, Haase, Matthaei. *JCAT* 1986; 10:363-8. (2) Helms, Dathe, Hagberg. *Concepts Magn Reson* 2004; **21A**: 50-62. (3) Helms, Piringner. *NMR Biomed* 2005; 18:44-50. (4) Natt, Watanabe, Boretius, Frahm, Michaelis. *Magn Reson Imag* 2003; 21:1113-20.

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