

# B<sub>1</sub>-Sensitivity Analysis of qMT

Mathieu Boudreau<sup>1</sup>, Nikola Stikov<sup>1</sup>, and G. Bruce Pike<sup>2</sup>

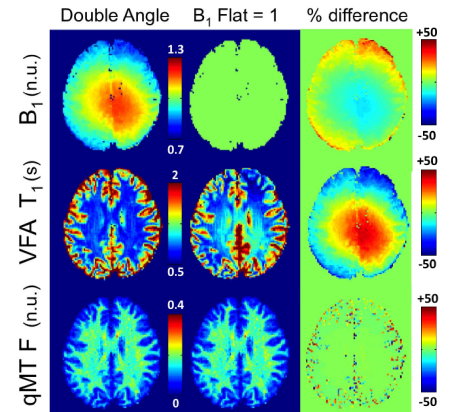
<sup>1</sup>McConnell Brain Imaging Center, Montreal Neurological Institute, McGill University, Montreal, Quebec, Canada, <sup>2</sup>Hotchkiss Brain Institute, Faculty of Medicine, University of Calgary, Calgary, Alberta, Canada

**INTRODUCTION:** B<sub>1</sub> mapping is an important measurement used in quantitative magnetization transfer (qMT) imaging, particularly at high field strengths ( $\geq 3.0$  T) where B<sub>1</sub> can vary by  $\pm 30\%$  in a human brain (Fig. 1). For pulsed spoiled gradient echo (SPGR) qMT imaging experiments, B<sub>1</sub> maps are used as a corrective factor for the excitation flip angle ( $\sim 5^\circ$  to  $15^\circ$ ) and MT saturation power (flip angles  $\sim 150^\circ$  to  $700^\circ$ ). Additional measurements necessary for qMT (e.g. T<sub>1</sub> mapping) may also require B<sub>1</sub> maps as a corrective factor; variable flip angle (VFA) T<sub>1</sub> mapping requires B<sub>1</sub> maps, while inversion recovery (IR) or Look-Locker typically do not<sup>1</sup>. Thus, local (e.g. artifacts) or global (e.g. systemic biases) inaccuracies in B<sub>1</sub> mapping<sup>2</sup> will propagate to the fitted qMT parameters differently, depending on the chosen T<sub>1</sub> mapping method.

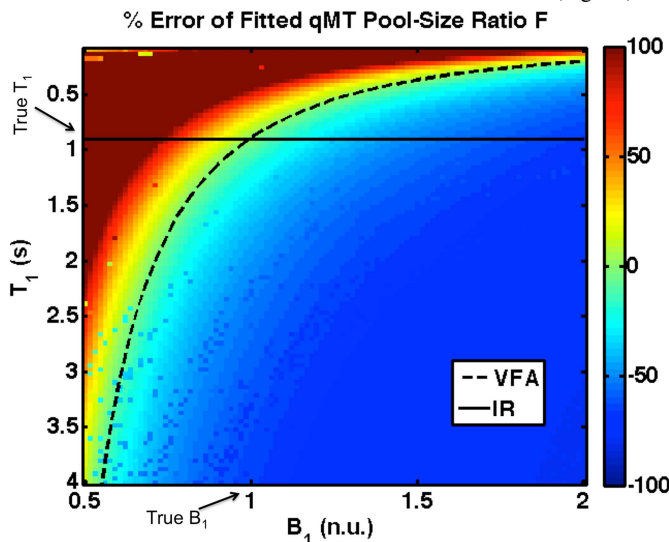
We recently reported that the qMT pool-size ratio (F), an important myelin biomarker, is insensitive to a large range of B<sub>1</sub> inaccuracies when using VFA for T<sub>1</sub> mapping<sup>3</sup> (Fig. 1). Here we present a simulation-based analysis of the B<sub>1</sub> sensitivity of qMT, comparing how different T<sub>1</sub> mapping methods (VFA vs. IR) propagate the B<sub>1</sub> error to the qMT parameters. We show that the F parameter is very robust and insensitive to B<sub>1</sub> inaccuracies when VFA T<sub>1</sub> mapping is used, but this comes at the expense of a substantial increase in error of kf.

**METHODS:** The Bloch-McConnell equations for magnetization exchange were solved using MATLAB (MATLAB2011a, The Mathworks Inc.) for a pulsed SPGR experiment by decomposing the pulse sequence into periods of instantaneous saturation of the free pool, constant irradiation of the restricted pool, and free precession<sup>4</sup>. Healthy white matter tissue parameters were fixed to the following values:  $F = 0.122$ ,  $k_f = 3.97 \text{ s}^{-1}$ ,  $R_{1f} = 1.11 \text{ s}^{-1}$ ,  $R_{1r} = 1.0 \text{ s}^{-1}$ ,  $T_{2f} = 27.2 \text{ ms}$ ,  $T_{2r} = 10.96 \mu\text{s}$ . The MT signal was simulated from the solution of the Bloch-McConnell equation for the following MT protocol: TR = 25 ms,  $\alpha_{\text{excitation}} = 7^\circ$ , Gaussian-Hanning MT pulses with a pulse duration of 10.2 ms,  $\alpha_{\text{MT}} = 142^\circ$  and  $426^\circ$ , logarithmically spaced off-resonance frequencies = 423.9 Hz, 1,087.5 Hz, 2,731.6 Hz, 6,861.6 Hz, and 17,235.4 Hz. The MT signal was subsequently fitted using the Sled and Pike method<sup>5</sup> for a linear range of 100 B<sub>1</sub> and 100 T<sub>1</sub> values (10,000 points in total); T<sub>1</sub> varied independently of B<sub>1</sub> for this step, and without any assumptions on the measurement method. B<sub>1</sub> ranged from 0.5 to 2 (B<sub>1,true</sub> = 1), and T<sub>1</sub> ranged from 0.1 s to 4 s (T<sub>1,true</sub> = 0.9 s). VFA signals were also simulated from the analytical SPGR equation<sup>1</sup>: TR = 25 ms,  $\alpha = 3^\circ$  and  $20^\circ$ , T<sub>1</sub> = T<sub>1,true</sub>. T<sub>1</sub> values were fitted from the VFA data for the B<sub>1</sub> error range. The fitted T<sub>1</sub> values were subsequently used in conjunction with their respective B<sub>1</sub> values to fit the MT signal.

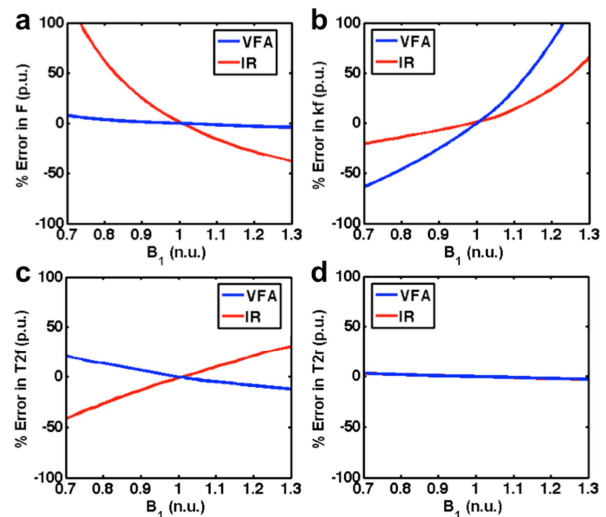
**RESULTS:** Figure 2 shows the error (%) of the fitted qMT pool-size ratio, F, in the presence of a wide range of B<sub>1</sub> and T<sub>1</sub> inaccuracies (B<sub>1,true</sub> = 1, T<sub>1,true</sub> = 0.9 s). The superimposed lines show the range of errors expected from an experiment using a B<sub>1</sub> independent T<sub>1</sub> method like IR (solid line), and from VFA T<sub>1</sub> mapping (dashed line). Figure 3 plots the errors in qMT fitted parameters (F, kf, T<sub>2f</sub>, T<sub>2r</sub>) using B<sub>1</sub>-independent (IR) and VFA measured T<sub>1</sub> (see lines in Fig. 2), for a range of B<sub>1</sub> inaccuracies typically observed in vivo. Errors in F induced by B<sub>1</sub> errors were greatly reduced using VFA T<sub>1</sub> mapping (Fig. 3a). A substantial increase in errors in kf occurs for VFA relative to IR (Fig. 3b), while T<sub>2r</sub> remains insensitive to B<sub>1</sub> inaccuracies for both cases.



**Figure 1.** Comparison of VFA T<sub>1</sub> and qMT F maps using measured and nominal (B<sub>1</sub> flat = 1) B<sub>1</sub> maps at 3T<sup>3</sup>.



**Figure 2.** Percent error in fitted qMT F values in the presence of a wide range of B<sub>1</sub> and T<sub>1</sub> errors (B<sub>1,true</sub> = 1 n.u., T<sub>1,true</sub> = 0.9 s). The superimposed lines plot the T<sub>1</sub> distribution for a B<sub>1</sub>-independent T<sub>1</sub> mapping method (IR, solid line) and VFA (dashed line).



**Figure 3.** Percent error in fitted qMT parameters for a range of B<sub>1</sub> errors (a – pool size ratio (F), b – magnetization exchange rate (kf), c – free pool T<sub>2</sub> (T<sub>2f</sub>), d – restricted pool T<sub>2</sub> (T<sub>2r</sub>)). Fits using a B<sub>1</sub>-independent T<sub>1</sub> measure (IR) are shown in red, and those using VFA T<sub>1</sub> mapping are shown in blue. See solid and dashed lines in Fig. 2 for B<sub>1</sub> dependence of IR and VFA T<sub>1</sub>.

**DISCUSSION:** The qMT pool size ratio F was shown to be nearly B<sub>1</sub>-error insensitive when using VFA T<sub>1</sub> mapping (Fig. 3a - blue). Using a B<sub>1</sub>-independent T<sub>1</sub> measure such as IR produces large qMT F errors (from  $>100\%$  to  $-45\%$  for B<sub>1</sub> errors ranging from  $-30\%$  to  $30\%$ , Fig 3a - red), while VFA T<sub>1</sub> mapping kept qMT F errors within a moderate range (7% to  $-3\%$ , Fig. 3a - blue). The B<sub>1</sub> errors for the case of VFA were mostly absorbed by the kf parameters (Fig. 3b), in agreement with observations from previous in vivo work<sup>3</sup>. These results suggest that qMT imaging using B<sub>1</sub>-independent T<sub>1</sub> measurement, and qMT methods that fixes qMT model parameters, may have increased sensitivity to B<sub>1</sub>-inaccuracies. However, for applications where kf may be the biomarker of interest (e.g. cartilage imaging<sup>6</sup>, systemic inflammation<sup>7</sup>), a B<sub>1</sub>-independent measure of T<sub>1</sub> may be preferred instead of the VFA method. Further analytical sensitivity analysis of the qMT equations for different qMT measurement protocols could help determine optimal qMT protocols for reduced B<sub>1</sub>-inaccuracy sensitivity.

**REFERENCES:** [1] Stikov, n. et al, MRM, doi: 10.1002/mrm.25135 (2014) [2] Boudreau, M. et al, Proc. of ISMRM, #3207 (2014) [3] Boudreau, M. et al, Proc. of ISMRM, #3167 (2014) [4] Sled J. and Pike G. B., JMR, 145:24-36 (2000) [5] Sled J. and Pike G. B., MRM, 46:923-931 (2001) [6] Stikov, N. et al, MRM, 66:725-734 (2011) [7] Harrison, N. et al, Biological Psychiatry, DOI: 10.1016/j.biopsych.2014.09.023 (2014)