## 10-Minute High-Resolution Whole-Brain T<sub>1</sub> Mapping: A Comparison of Three Candidate Methods

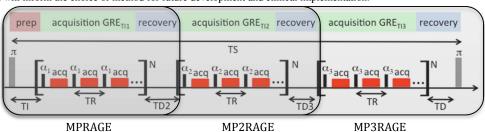
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Target Audience: Researchers and clinicians interested in fast, high-resolution T<sub>1</sub> mapping of whole organs (e.g. whole-brain), particularly at high field strengths.

**Purpose**: Accurate  $T_1$  maps can now be obtained with three MPRAGE images with suitably chosen inversion times  $(TI)^1$ , or two MP2RAGE images obtained during a single interleaved acquisition. <sup>2,3</sup> Such methods could enable high-resolution  $T_1$  mapping in clinically relevant scan times, i.e. on the order of 10 minutes, allowing for better visualization and classification of deep-brain structures. Due to their robustness to  $B_1$  heterogeneity, these MPRAGE-based methods are particularly attractive for high field applications. In this work, we compare three candidate sequences at 7T: 3-TI MPRAGE as described by Liu *et al.* <sup>1</sup>, MP2RAGE as described by Marques *et al.* <sup>2,3</sup>, and MP3RAGE, a modification of MP2RAGE first proposed by Hung *et al.* <sup>4</sup> that allows collection of a third image without increasing scan time. This experimental comparison of  $T_1$  accuracy and precision will inform the choice of method for future development and clinical implementation.

Figure 1. Pulse sequence schematic for MPRAGE-based  $T_1$  mapping methods. MPRAGE acquires an image segment at time TI after inversion with a train of N pulses of flip angle  $\alpha_i$ . 3-TI MPRAGE requires three acquisitions with varying TI but constant TS. MP2RAGE re-acquires the same segment before the next inversion pulse, after a delay TD2 (effective inversion time TI2=TI+N\*TR+TD2). MP3RAGE adds a third acquisition of the segment (effective inversion time TI3=TI2+N\*TR+TD3.



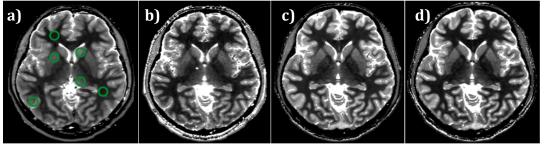
**Methods**: One to three images are acquired following each inversion pulse, as shown in Figure 1, with each acquisition collecting the same single k-space segment during the TS interval. For the 3-TI MPRAGE method, TS is fixed by altering the final delay TD for each TI; this ensures that all three images have a simple dependence on  $T_1$  that allows a straightforward lookup instead of a least-squares fit, and which removes any dependence on  $M_0$ ,  $T_2*$  or  $B_1$  effects. A similar lookup is used for MP2RAGE and MP3RAGE using appropriate combinations of the acquired images. These lookups were done without additional  $B_1$  correction, and in forming the lookup table the inversion pulse efficiency was set to -0.85 (empirically determined to produce best accuracy). All methods were implemented with a 2D centric (radial fan-beam<sup>5</sup>) phase encode ordering that reduces scan time by omitting k-space corners, allows for flexible  $k_y$ - $k_z$  undersampling, and also improves  $B_1$  insensitivity in the MP2RAGE and MP3RAGE images (data not shown).

| Parameter  | 3-TI MPRAGE     | MP2RAGE   | MP3RAGE       |
|------------|-----------------|-----------|---------------|
| rarameter  | 3-11 MITKAGE    | MPZKAGE   | WIFSKAGE      |
| TS (ms)    | 6000            | 7500      | 7500          |
| TI (ms)    | 150/1280/4000   | 1000/3300 | 700/2200/4000 |
| α (deg)    | 5               | 5/4       | 5/4/5         |
| N          | 240             | 200       | 200           |
| TR (ms)    | 7.7             | 7.7       | 7.4           |
| BW (kHz)   | 25              | 25        | 27.78         |
| ARC factor | 3x2             | 2x1       | 2x1           |
| Scan time  | 3:18 x 3 = 9:54 | 10:00     | 10:00         |

Table 1. Sequence parameters for candidate methods.

All images were obtained using a GE Discovery MR950 7T scanner (GE Healthcare, Waukesha WI) with a Nova 2chTx / 32chRx head coil (Nova Medical, Wilmington, MA). Two healthy volunteers (1M/1F, age 33 years) were scanned according to IRB requirements. Parameters for the MPRAGE-based methods are listed in Table 1. A reference  $T_1$  map was acquired with a single-slice IR-FSE sequence, ETL=8, TR=6000ms, TI=50, 200, 600, 1500 and 4000ms, with  $T_1$  determined by a reduced-dimension non-linear least-squares fit.<sup>6</sup> Scan time was kept constant at 10min for the three different MPRAGE methods and for the IR-FSE reference, with 1mm isotropic resolution in all cases.

**Results:** The single slice reference T<sub>1</sub> map from one volunteer is shown in Fig. 2a, along with T<sub>1</sub> maps from the corresponding slice of the whole-brain 3-TI MPRAGE (Fig. 2b), MP2RAGE (Fig. 2c) and MP3RAGE (Fig. 2d) scans. ROIs used for comparison are highlighted in green. The overall quality of the T<sub>1</sub> maps is similar, with MP2RAGE having an overall T<sub>1</sub>-to-Noise-Ratio (T<sub>1</sub>NR) advantage, which we calculate as the average of (ROI mean)/(ROI std.dev.) across all ROIs (see Table 2). In terms of accuracy compared to the reference map, computed as the average absolute % error from the reference, 3-TI MPRAGE has the best performance, followed by MP3RAGE.



| Method   | T <sub>1</sub> NR | %Error    |
|----------|-------------------|-----------|
| IR-FSE   | 30 +/- 10         | N/A       |
| 3xMPRAGE | 16 +/- 5          | 1.3 +/- 1 |
| MP2RAGE  | 32 +/- 16         | 4 +/- 2   |
| MP3RAGE  | 24 +/- 10         | 2 +/- 1.7 |

Figure 2. T<sub>1</sub> maps produced by (a) IR-FSE single-slice reference, (b) 3-TI MPRAGE, (c) MP2RAGE, (d) MP3RAGE.

**Discussion and Conclusions:** MP2RAGE has the highest  $T_1NR$  of the three methods compared, but the lowest accuracy, likely due to residual  $B_1$  effects. A separate  $B_1$  correction is possible but this adds to scan time and reconstruction complexity. The reduced  $T_1NR$  of 3-TI MPRAGE is caused by the shorter segment time and increased undersampling necessary to achieve a 10-minute scan (since MP2RAGE and MP3RAGE acquire all images within a single TS, less acceleration is needed for the same scan time). Our implementation of MP3RAGE is a promising compromise that offers higher  $T_1NR$  than 3-TI and lower relative error than MP2RAGE.

**References:** [1] Liu et.al., *NeuroImage* **56** pp.1154–1163 (2011) [2] Marques et.al., *NeuroImage* 49 (2010). [3] Marques and Gruetter, *PLoS ONE* 8:7 (2013). [4] Hung et.al., Proc. ISMRM #2353 (2013). [5] Saranathan and Glockner, *JMRI* doi:10.1002/jmri.24113 (2013). [6] Barral et.al., *MRM* **64** pp.1057-1067 (2010).

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