## Multiple Echo and Inversion Time MPRAGE with Inner Loop GRAPPA Acceleration and Prospective Motion Correction for Minimally Distorted Multispectral Brain Morphometry

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Target audience: Clinicians and researchers interested in efficient high quality brain imaging, multispectral morphometry and tissue parameter mapping with a single modified MPRAGE acquisition.

Purpose: Distortions and blurring in MPRAGE affect morphometric measurements (e.g. cortical thickness) that are signatures of brain disease<sup>1</sup>.

Remaining B<sub>0</sub>-inhomogeneities after shimming result in accumulated phase errors across each readout and therefore erroneous spatial encoding (susceptibility distortions). In addition, T<sub>2</sub>\*-decay across the readout results in broadening of the point spread function (PSF). Replacing the single, low-bandwidth gradient echo readout with multiple shorter, highbandwidth readouts results in reduced susceptibility distortion and a narrower PSF in the readout direction. SNR is recovered by combining the readouts in image reconstruction. T2\* may be estimated by fitting signal decay across the echoes.

T<sub>1</sub>-recovery during partition (k-space "slice") encoding after each inversion (MP) pulse in 3D MPRAGE similarly results in a broadened PSF in this phase encoding direction. Partition encoding time can be reduced by GRAPPA acceleration in this inner phase encoding loop ("inner-loop GRAPPA", ILG), thus reducing distortion but also reducing SNR. Replacing each partition encoding block with multiple ILG-accelerated blocks experiencing different inversion times recovers SNR when the partitions are combined in image reconstruction. T<sub>1</sub> may then be estimated by fitting the signal predicted by Bloch simulation across inversion times<sup>2</sup>

Images may be blurred further by subject motion during the long high resolution acquisitions, which may require several minutes even with outer loop acceleration.

Methods: We implemented a generalized MPRAGE sequence<sup>2</sup> that allows ≤12 gradient echoes, ≤8 inversion times, ILG acceleration, and incorporates vNavs for real-time prospective motion correction and reacquisition of damaged TR intervals<sup>3</sup>. The excitation scheme is shown in Figure 1 alongside that of a standard MPRAGE protocol recommended for brain morphometry4. Motion-corrected volumes for each TE and TI are reconstructed immediately on the scanner using improved "IcePat" routines for GRAPPA acceleration.

The method was evaluated in two volunteers on a 3 T Skyra (Siemens Healthcare, Erlangen) with a 32-channel head coil. The protocol was selected to match the timing of the conventional 1x1x1 mm³ 3D MPRAGE (no acceleration), with optimal gray matter/white matter/CSF contrast for morphometry⁴. MEMPxRAGE parameters: 2 gradient echoes, 3 inversion times, TR 2.53 s, TE 1.69/3.55 ms, BW 650 Hz/px, TI 700/1400/2100 ms, flip angle matter at 3 T, sampled at times of ADC reads. 7/7/7°, 3x ILG acceleration (40 ref. lines), 176 sagittal slices, 256x256 matrix, 1x1x1 mm resolution, Tacq 11:21 min:s. vNav parameters: 3D EPI, TR 11 ms, TE 5.1 ms, 32x32 matrix, 32 sagittal slices,

6/8 partition partial Fourier, 8×8×8 mm<sup>3</sup> voxels, BW 4464 Hz/px, T<sub>acq</sub>/nav 275 ms.

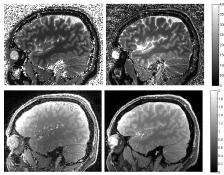


Figure 3: Estimates of T<sub>1</sub> (top) and PD (bottom), MEMPxRAGE-ILG acquisition/Bloch (left) and multiecho FLASH acquisition/model equation fit (right).

A forward model for signal generation based on Bloch equations was implemented in Matlab (MathWorks, Natick, MA) and used to predict signals for a given T<sub>1</sub> relaxation time<sup>2</sup> (Figure 1). Figure 2 shows corresponding PSFs for each sequence. Note the broader PSF for conventional MPRAGE. The proton density (PD) and T<sub>1</sub> time that best predicted the observed signal evolution was found at each voxel across the range of inversion times. For comparison, two multiecho FLASH (MEF) volumes were acquired on the same subjects: TR 20 ms, TE 1.91+1.9*n*, *n*=0,...,7, BW 650 Hz/px, flip angles  $5^{\circ}$  and  $30^{\circ}$  (separate acquisitions),  $T_{acq}$ 

inner-phase encoding direction for (left) MEMPxRAGE-ILG TI 1 (blue), 2 (green) and 3 (red) and (right) MPRAGE. Profiles scaled by mean modeled signal intensity. 5:08 min:s, 4× GRAPPA acceleration, geometry matches scaled by mean modeled signal in MEMPxRAGE-ILG. Standard methods<sup>5,6</sup> were used to estimate PD and T<sub>1</sub> from the MEF scans. Results: Figure 3 shows T1 and PD estimates from MEMPxRAGE-ILG using Bloch simulation, and estimates for the same subject derived by fitting the steady-state equation for the MEF scans<sup>5,6</sup>. Figure

4 shows a FreeSurfer surface model/parcellation from the combined MEMPxRAGE-ILG volumes. Discussion: GRAPPA successfully recovers volumes at all TIs and TEs with minimal residual aliasing.

The more uniform signal over the partition at each inversion time (due to acceleration) together with

the availability of more than two inversion times better constrains the Bloch simulation and may provide better T<sub>1</sub> fits. Results are comparable to T<sub>1</sub> fitting with MEF, although the latter may be more affected by dielectric resonance effects. The effect of the embedded vNavs is subtle and can be modeled (Figure 1). As in traditional FLASH acquisitions and with appropriate timing, the multiple gradient echoes from the MEMPxRAGE-ILG sequence can be used to estimate the B<sub>0</sub> field map, separate fat from water, and estimate T<sub>2</sub>\* decay at each voxel using traditional techniques<sup>7,8</sup> Bloch simulation can also be used to fit T<sub>2</sub>\*, however more than two echoes would be preferable. Conclusion: MEMPxRAGE with ILG and vNavs is an efficient sequence for high quality, low

Figure 2: PSF for white matter at 3 T in

Figure 4: Surface reconstruction (left) and surface parcellation (right) from MEMPxRAGE-ILG.

distortion and low blur acquisitions, useful for quantitative tissue parameter mapping and high resolution brain morphometry in a single acquisition. Acknowledgements: This work was supported by: NIH R21MH096559, R01HD071664, R21EB008547, K01EB011498, K99HD074649, P41RR014075, and the Ellison Medical Foundation. Thanks to Drs. T. Kober and G. Krueger for access to image sorting code.

References: [1] Bakkour et al., Neurology 72(12):1048-55, 2009. [2] Marques et al., Neuroimage 49(2):1271-81, 2010. [3] Tisdall et al., MRM 68(2):389-99, 2012. [4] Van der Kouwe et al., Neuroimage 40(2):559-69, 2008. [5] Deoni et al., MRM 53(1):237-41, 2005. [6] Fischl et al., Neuroimage 23 suppl 1:S69-84, 2004. [7] Glover, JMRI 1(5):521-30. [8] Yu et al., JMRI 26(4):1153-61.

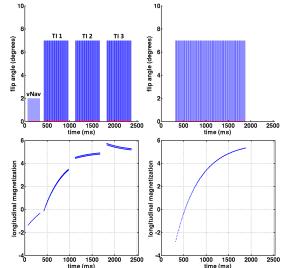


Figure 1: Single steady-state TR (between inversions) of (right) conventional (left) MEMPxRAGE-ILG and MPRAGE sequence. (Top) Excitation scheme (blue), ADC events (red). (Bottom) Signal evolution for white