PAPER • OPEN ACCESS

Evaluation of MRI sequences for quantitative T1 brain mapping

View the article online for updates and enhancements.

Evaluation of MRI sequences for quantitative T1 brain mapping

P Tsialios[1], M Thrippleton[2], A Glatz[2] and C Pernet[2]

[1] Department of Biomedical Engineering, Technological Educational Institute of Athens

[2] Centre for Clinical Brain Sciences, University of Edinburgh

E-mail: ptsialios@teiath.gr

Abstract. T1 mapping constitutes a quantitative MRI technique finding significant application in brain imaging. It allows evaluation of contrast uptake, blood perfusion, volume, providing a more specific biomarker of disease progression compared to conventional T1-weighted images. While there are many techniques for T1-mapping there is a wide range of reported T1-values in tissues, raising the issue of protocols reproducibility and standardization. The gold standard for obtaining T1-maps is based on acquiring IR-SE sequence. Widely used alternative sequences are IR-SE-EPI, VFA (DESPOT), DESPOT-HIFI and MP2RAGE that speed up scanning and fitting procedures. A custom MRI phantom was used to assess the reproducibility and accuracy of the different methods. All scans were performed using a 3T Siemens Prisma scanner. The acquired data processed using two different codes. The main difference was observed for VFA (DESPOT) which grossly overestimated T1 relaxation time by 214 ms [126 270] compared to the IR-SE sequence. MP2RAGE and DESPOT-HIFI sequences gave slightly shorter time than IR-SE (~20 to 30ms) and can be considered as alternative and time-efficient methods for acquiring accurate T1 maps of the human brain, while IR-SE-EPI gave identical result, at a cost of a lower image quality.

1. Introduction

Among clinical examination modalities, Magnetic Resonance Imaging (MRI) is one of the most widely used, allowing to distinguish pathologic tissues with great precision. Contrast images like T1-weighted images are for instance used routinely as they preserve well anatomical details. These types of images, however, are qualitative images in the sense that they do not accurately measure tissue parameters such as T1 recovery time. Research efforts have shown interest in quantifying the longitudinal relaxation time (T1), especially in brain tissue [1] because of an increased sensitivity in pathology detection such as brain cancer, multiple sclerosis (integrity of myelin in the brain), ischemia, hepatic encephalopathy, or chronic alcoholism [2-5]. Moreover, quantitative T1 maps can help clinicians to evaluate contrast agent uptake, iron overload, and blood perfusion and volume [6]. They also provide a more robust template for morphometry studies and a more specific marker of disease progression in comparison to conventional T1 weighted images [7].

The gold standard method for T1 mapping makes use of inversion recovery spin echo (IR-SE) sequence [8,9]. However, IR-SE requires an extremely long repetition time (TR) resulting in long scan time, which is unpractical for clinical use and can lead to inaccurate measurements due to patient motion. Many efforts have been made to shorten the T1 measurement time, leading to a variety of techniques and a wide range of reported T1 values (from ~1000 to 1600ms) raising the issue of protocols reproducibility and standardization [10]. The purpose of this study is to estimate the reproducibility and accuracy of T1 values acquired by four alternative T1 mapping protocols in a phantom and compare them to T1 values acquired by a gold standard protocol [6].

2. Methods

2.1 Imaging protocols

All imaging was performed at the University of Edinburgh on a 3Tesla Siemens Prisma scanner using a custom MRI phantom [11]. The phantom (figure 1a) is filled with 1.5 g/l $CuSO_4$ and 3.6 g/l NaCl solution and made of 9 tubes containing $MnCl_2$ solution of various concentrations. Five different pulse sequences were used to image the phantom: a) IR-SE (TI = 30, 330 530, 780, 1030, 1530ms, TE/TR = 11/1550ms), b) IR-SE Echo Planar Imaging (IR-SE-EPI) (TI = 100, 340, 580, 820, 1060, 1300, 2000, 3000ms, TE/TR = 42/20000ms), c) Inversion Recovery Spoiled Gradient Recalled Acquisition in Steady State (IR-SPGR) or DESPOT-HIFI (TI = 600, 1500ms, TE/TR = 1,91/1170ms, flip angles = 2°, 5°, 12°), d) Variable Flip Angle (VFA) spoiled gradient recalled echo (flip angles = 2°, 5°, 12°) and e) Three Dimensional Magnetization Prepared Rapid Acquisition GRE (MP2RAGE) (TE/TR = 2.98/5000ms). A thorough recording of phantom and room temperature was conducted in order to ensure the same experimental conditions as T1 increases with temperature by 2–3% per degree Celsius [12].

2.2 Image Processing

Image processing was carried out using two Matlab codes, the Stanford code (part of theT1Mapping Matlab package which was developed by Nikola Stikov and his counterparts at Stanford University [6]) and an in-house code (MJT) that uses a different method to fit the data. Both codes accept DICOM images where magnitude, phase, real part and imaginary part have been saved. Reported T1 values were obtained from each test tube by taking the average values across voxels for each TR (except MP2RAGE that provides directly the T1 value). Statistical comparisons were performed using a percentile bootstrap on median differences, with adjustment for multiple comparisons [13].

3   Results

The reproducibility of the gold standard IR-SE was evaluated by acquiring the sequence for four consecutive days (day1: 866ms [531 1143], day2: 748ms [509 1016] day3: 745ms [511 1023] day4: 786ms [564 1070], figure 1b). Small but significant fluctuations of T1 values were observed (p=.001) that might have risen from distributions in phantom temperature. Code comparison was performed on IR-SE and IR-SE-EPI sequences. Fitting procedures did not show difference on T1 values (2ms [-6 13] and 4ms [-9 9] respectively). Given the above results, comparison of sequences was performed from data acquired the same day and using MJT code (table 1). Significant differences were observed with VFA (DESPOT) showing much longer T1 values than other sequences (p=.001, +228ms on average). DESPOT-HIFI (-11ms [-22 -4] p=.002) and MP2RAGE (-32ms [-45 -25] p=.001) had shorted values than the gold standard, with a significant difference between them (p=.001) (figure 1c and 1d).

Table 1. T1 values in ms (with median and 95% highest density interval) in each ROI.

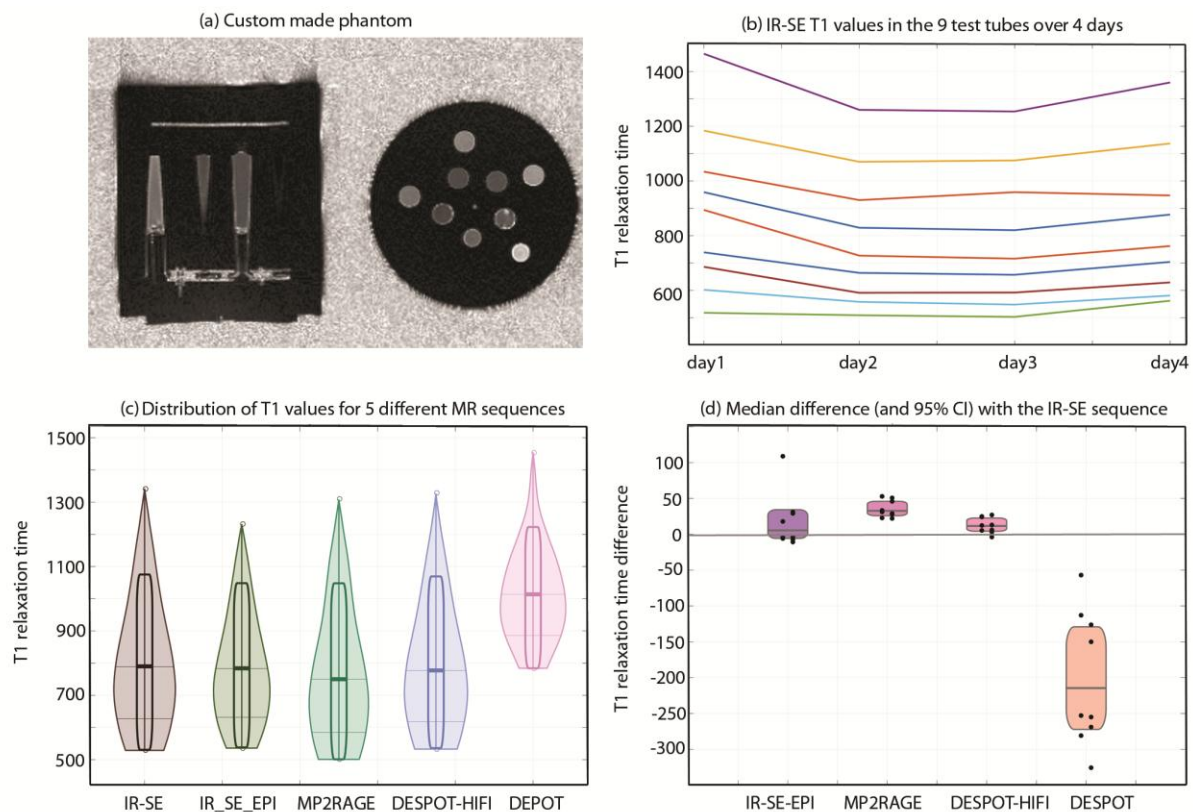| ROI	IR-SE	IR-SE-EPI	M2RAGE	DESPOT-HIFI	VFA(DESPOT)
1	872	843	850	866	1022
2	987	969	964	962	1044
3	1131	1100	1102	1104	1257
4	1341	1232	1310	1329	1454
5	529	536	501	533	784
6	582	588	549	577	835
7	631	642	585	618	900
8	702	706	649	678	983
9	763	767	712	760	1089
median	790	783	750	777	1013
HDI	[530 1075]	[538 1048]	[501 1048]	[535 1069]	[784 1222]

Figure 1. Panel (a) shows the MP2RAGE image of the phantom, panel (b) shows the T1 values acquired with IR-SE over 4 consecutive days, panel (c) shows the distributions of T1 values for IR-SE, IR-SE-EPI, MP2RAGE, DESPOT-HIFI and VFA (DESPOT) acquired on the same day and panel (d), depicts the relative differences from IR-SE-EPI, MP2RAGE, DESPOT-HIFI and VFA (DESPOT) to the gold standard IR-SE sequence. Violin plots show the non-parametric kernel of each data (Random Average Shifted Histograms) and rectangles indicate the 95% confidence intervals of the medians (thick lines).

4 Discussion and Conclusion

The advantages of using quantitative imaging and especially T1 mapping methods in MRI studies are evident. They should provide unambiguous evidence for the change of various pathologic conditions. However, the main difficulty in current T1 mapping protocols is the impractically long scan time creating the need for new faster ones. Our results show that the gold standard IR-SE is relatively stable but some values seems more sensitive to temperature variations than others, b) data fitting procedure have little impact on T1 values c) VFA (DESPOT) T1 maps give longer T1 values in comparison to the other T1 maps, d) MP2RAGE and DESPOT-HIFI T1 values are close to the gold standard but still differ significantly.

References

[1] Fatouros P and Marmarou A 1999 Use of magnetic resonance imaging for in vivo measurements of water content in human brain: method and normal values *J Neurosurg* **90 (1)** 109-15

[2] Vrenken H *et al*. 2006 Whole-Brain T1 Mapping in Multiple Sclerosis: Global Changes of Normal-appearing Gray and White Matter *Radiology* **240 (3)** 811-20

[3] Neema M, Stankiewicz J, Arora A, Dandamudi V, Batt C, Guss Z, Al-Sabbagh A and Bakshi R 2007 T1- and T2-Based MRI Measures of Diffuse Gray Matter and White Matter Damage in Patients with Multiple Sclerosis *Neuroimaging* **17 (Supplement s1)** 16S-21S

[4] Shah N, Neeb H, Zaitsev M, Steinhoff S, Kircheis G, Amunts K., Häussinger D and Zilles K 2003 Quantitative T1 mapping of hepatic encephalopathy using magnetic resonance imaging *Hepatology* **38 (5)** 1219-26

[5] Steen R, Hunte M, Traipe E, Hurh P, Wu S, Bilaniuk L and Haselgrove J 2004 Brain T1 in young children with sickle cell disease: evidence of early abnormalities in brain development *Magn Reson Imaging* **22 (3)** 299-06

[6] Barral J, Gudmundson E, Stikov N, Etezadi-Amoli M, Stoica P and Nishimura D 2010 A Robust Methodology for In Vivo T1 Mapping *Magn Reson Med* **64 (4)** 1057-67

[7] Lorio S, Ferath K, Ruef A, Melie-Garcia L, Frackowiak R, Ashburner J, Helms G, Lutti A and Draganski B 2016 Neurobiological origin of spurious brain morphological changes: A quantitative MRI study *Hum Brain Mapp* **37(5)** 1801-15

[8] Drain L 1949 A Direct Method of Measuring Nuclear Spin-Lattice Relaxation Times *Proceedings of the Physical Society* Section A **62 (5)** 301-06

[9] Hahn E 1949 An Accurate Nuclear Magnetic Resonance Method for Measuring Spin-Lattice Relaxation Times *Phys. Rev.* **76** 145-46

[10] Kingsley P 1999 Methods of measuring spin-lattice (T1) relaxation times: an annotated bibliography *Concepts Magn Reson* **11** 243–76

[11] Glatz A, Bastin M, Kiker A, Deary I, Wardlaw J and Valdes Hernandez M 2015 Automated segmentation of multifocal basal ganglia T2*-weighted MRI hypoitensities *Neuroimage* **105** 332-46

[12] Nelson T and Tung S 1987 Temperature dependence of proton relaxation times in vitro *Magn Reson Imaging* **5** 189–99

[13] Wilcox R 2012 *Introduction to Robust Estimation and Hypothesis Testing 3rd Edition* Elsevier Academic Press (San Diego CA USA) chapter 8 p398-01