# **Reproducibility of Myocardial T<sub>1</sub> and T<sub>2</sub> Relaxation Time**

# Measurement using Slice-Interleaved T<sub>1</sub> and T<sub>2</sub> Mapping Sequences

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# Abstract

**Purpose:** To assess measurement reproducibility and image quality of myocardial  $T_1$  and  $T_2$  maps using free-breathing slice-interleaved  $T_1$  and  $T_2$  mapping sequences at 1.5T.

**Material and Methods:** Eleven healthy subjects ( $33\pm16$  years, 6 males) underwent a sliceinterleaved T<sub>1</sub> and T<sub>2</sub> mapping test/retest cardiac MR study at 1.5T on 2 days. For each day, subjects were imaged in two sessions with removal out of the magnet and repositioning prior to the subsequent session. We studied measurement reproducibility as well as the required sample size for sufficient statistical power to detect a predefined change in T<sub>1</sub> and T<sub>2</sub>. In a separate prospective study, we assessed T<sub>1</sub> and T<sub>2</sub> map image quality in 241 patients ( $54 \pm 15$  years, 73 women) with known/suspected cardiovascular disease referred for clinical cardiac MR. A subjective quality score was used to assess a segment-based image quality.

**Results:** In the healthy cohort, the slice-interleaved  $T_1$  measurements were highly reproducible, with global coefficients of variation (CV's) of 2.4% between subjects, 2.1% between days and 1.7% between sessions. Slice-interleaved  $T_2$  mapping sequences provided similar reproducibility with global CV's of 7.2% between subjects, 6.3% between days and 5.0 between sessions. A lower variability resulted in a reduction of the required number of subjects to achieve a certain statistical power when compared to other  $T_1$  mapping sequences. In the subjective image quality assessment, >80% of myocardial segments had interpretable data.

**Conclusion:** Slice-interleaved  $T_1$  and  $T_2$  mapping sequences yield highly reproducible  $T_1$  and  $T_2$  measurements with >80% of interpretable myocardial segments.

**Keywords:** Myocardial tissue characterization and relaxometry, sample size justification, sliceinterleaved  $T_1$  and  $T_2$  mapping sequence.

# **INTRODUCTION**

Myocardial interstitial diffuse fibrosis, inflammation and edema are present in many diseases such as cardiomyopathy (1), hypertension (2), aortic regurgitation (3), myocarditis (4). Therefore, non-invasive assessment of myocardial tissue composition of fibrosis, edema and inflammation may have an important clinical impact in diagnosis, prognosis and monitoring of therapy. Myocardial tissue relaxometry has been emerging as a clinically powerful tool to characterize myocardial tissue composition (5). Myocardial T<sub>1</sub> relaxation time changes in the presence of interstitial diffuse fibrosis and can be assessed using native T<sub>1</sub> and extra-cellular volume (ECV) mapping (6,7). Changes in myocardial T<sub>2</sub> relaxation time in the presence of edema and inflammation can also be quantitatively measured using myocardial T<sub>2</sub> mapping (4,8,9).

Over the past decade, there have been numerous advances in cardiac MR sequences for myocardial  $T_1$  and  $T_2$  mapping (10-23).  $T_1$  mapping sequences sample the magnetization recovery curve to enable estimation of  $T_1$  recovery by voxel-wise curve-fitting. The modified Look-Locker inversion recovery (MOLLI) (13) and shortened MOLLI (ShMOLLI) (14) sequences are widely used for myocardial  $T_1$  mapping. Saturation recovery based sequences such as modified Look-Locker acquisition using saturation recovery (15), SAturation recovery single-SHot Acquisition (SASHA) (16), and saturation method using adaptive recovery times for cardiac  $T_1$  mapping (SMART<sub>1</sub>Map) sequence (17) have also been investigated. These sequences allow more accurate  $T_1$  measurements; however, with a penalty in precision and reproducibility (18). A combination saturation and inversion recovery (SAPPHIRE) (19) sequence has also been investigated. A free-breathing slice-interleaved  $T_1$  mapping sequence (STONE) (20) has been

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recently proposed that enables  $T_1$  measurements of 5 short-axis slices over the entire left ventricle in a free-breathing 95 second scan. This sequence is based on interleaving data acquisition for different slices during the recovery time of adjacent slices. A longer recovery of spins for each individual slice results in improved accuracy and precision (20).

Myocardial T<sub>2</sub> mapping is often performed by acquiring multiple T<sub>2</sub> prepared (T<sub>2</sub>prep) balanced steady-state free-precession images, each followed by rest periods of magnetization recovery, and estimating the voxel-wise T<sub>2</sub> values (21). This is often performed with one breath-hold for each slice. 3D myocardial T<sub>2</sub> mapping is an alternative to 2D T<sub>2</sub> mapping so as to improve spatial resolution and coverage of myocardial T<sub>2</sub> mapping (22). A slice-interleaved T<sub>2</sub> mapping sequence was recently developed by implementing a slice-selective T<sub>2</sub>Prep to interleave the data acquisition for different slices in subsequent heartbeats (23). This free-breathing slice-interleaved T<sub>2</sub> mapping sequence allows T<sub>2</sub> measurements of five parallel left ventricular short-axis slices with similar precision as a single-slice T<sub>2</sub> mapping sequence, but with a 4-fold reduction in acquisition time (23).

In recent years, there has been interest in applying regional myocardial  $T_1$  and  $T_2$  mapping as imaging markers of disease progression or response to a specific therapy/intervention. Knowledge of measurement reproducibility is important to distinguish between changes that could be attributable to measurement variability and those that are the result of disease progression or therapeutic intervention. Furthermore, to justify the sample size for the achievement of a specific statistical power, the reproducibility of these measurements needs to be known. Finally, image quality assessment should be performed to take into account the percentage of non-diagnostic images acquired with these sequences. To address these challenges, we sought to: 1) investigate reproducibility of recently developed free-breathing slice-interleaved

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 $T_1$  and  $T_2$  mapping sequences in a cohort of healthy adult subjects by performing a comprehensive test/re-test study; 2) investigate the required sample size to achieve certain statistical power for detection of a pre-defined change in native  $T_1$  or  $T_2$ ; and 3) assess overall  $T_1$  and  $T_2$  map image quality using the slice-interleaved  $T_1$  and  $T_2$  mapping sequences in patients with known or suspected cardiovascular disease referred for clinical cardiac MR.

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# **MATERIAL AND METHODS**

All imaging was performed on a 1.5T Philips Achieva (Philips Healthcare, Best, The Netherlands) MRI system using a 32-channel cardiac coil. The study was Health Insurance Portability and Accountability Act (HIPAA) compliant. The imaging protocol was approved by our institutional review board and written informed consent was obtained from each participant prior to examination.

# **Reproducibility Assessment**

In a prospective study, we recruited 11 healthy adult subjects ( $33\pm16$  years; 6 men) without contraindications for cardiac MR to participate in a 2-day test/retest study using an imaging protocol shown in **Figure 1**. Each subject underwent cardiac MR imaging on two different days (between-day reproducibility) with the identical imaging protocol. After image localization, the subjects were imaged using slice-interleaved T<sub>1</sub> and T<sub>2</sub> mapping sequences in five left ventricular short-axis view slices over the entire ventricle (from apex to base). Each sequence was repeated twice (within-session reproducibility). Following completion of the first imaging session, subjects were taken out of the scanner for a 10-15 minutes break with removal of the coil. Subsequently, subjects were scanned again - after image localization - for a second session (between-session reproducibility) with two repetitions per sequence.

T<sub>1</sub> mapping was performed using slice-interleaved T<sub>1</sub> mapping sequence with the following parameters: 5 short-axis slices, in-plane resolution=  $2.1 \times 2.1 \text{ mm}^2$ , slice thickness=8 mm, slice gap=4 mm, field-of-view= $320 \times 320 \text{ mm}^2$ , TR/TE/ $\alpha$ =2.8 ms / 1.38 ms / 70°, SENSE-rate=2, linear ordering, 10 linear ramp-up pulses and acquisition window=218.8 ms, bandwidth=1879.7

Hz/pixel. T<sub>2</sub> mapping was performed using slice-interleaved T<sub>2</sub> mapping sequence with the following parameters: 5 short-axis slices, in-plane resolution= $2.1 \times 2.1 \text{ mm}^2$ , slice thickness=8 mm, slice gap=4 mm, field-of-view= $320 \times 320 \text{ mm}^2$ , TR/TE/ $\alpha$ = 2.8 ms / 1.38 ms / 55°, SENSE-rate=2.3, linear ordering, 10 linear ramp-up pulses and acquisition window=191.1ms, bandwidth=1879 Hz/pixel. Images for both T<sub>1</sub> and T<sub>2</sub> mapping were acquired during free-breathing with slice tracking. A two-dimensional pencil beam navigator was positioned at the dome of the right hemi-diaphragm to monitor the diaphragmatic motion and to correct the slice position during imaging. Only slice tracking was used without any respiratory gating.

### **Impacts on Sample Size Calculation**

Using  $T_1$  and  $T_2$  measurements from our healthy subject cohort, we performed statistical analyses to determine the number of subjects that are needed to achieve a certain power for detection of specific changes in  $T_1$  and  $T_2$ . For sample size assessment, we compared the results of slice-interleaved  $T_1$  with  $T_1$  measured using MOLLI and ShMOLLI. As we did not directly acquire data using MOLLI or ShMOLLI, this variability was extracted from a previously published study (18). Correspondingly, we compared the results of slice-interleaved  $T_2$  with  $T_2$  measured by using single-slice  $T_2$  mapping sequence. The data for the single-slice  $T_2$  sequence was acquired in the same sessions as the slice-interleaved sequences with our 11 healthy subjects using the identical study design. The sequence was performed by using the following parameters: 5 short-axis slices, in-plane resolution= $2.1 \times 2.1 \text{ mm}^2$ , slice thickness=8 mm, slice gap=4 mm, field-of-view= $320 \times 320 \text{ mm}^2$ , TR/TE/ $\alpha$ = $2.9 \text{ ms} / 1.43 \text{ ms} / 55^\circ$ , SENSE-rate=2.3, linear ordering, 10 linear ramp-up pulses and acquisition window=197.3 ms, bandwidth=1879 Hz/pixel.  $T_1$  and

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 $T_2$  measurements from the mid LV slice were used for all calculations in the sample size analysis.

# **Image Quality Assessment**

For patient image quality assessment, we prospectively enrolled 246 patients (54±15 years, 73 women) with known or suspected cardiovascular disease referred for a clinical cardiac MR exam over a period of 15 months. The imaging protocol was approved by our institutional review board and written informed consent was obtained from each participant prior to each examination for the addition of  $T_1$  and  $T_2$  mapping sequences to their standard clinical exam. Slice-interleaved T<sub>1</sub> and T<sub>2</sub> mapping sequences were acquired in addition to their clinically indicated imaging protocol. Imaging parameters were similar to the healthy subject study. The slice-interleaved T<sub>1</sub> mapping sequence had the following imaging parameters: 5 short-axis slices, in-plane resolution= $2.1 \times 2.1 \text{ mm}^2$ , slice thickness=8 mm, slice gap=4 mm, field-ofview=360×352 mm<sup>2</sup>, TR/TE/ $\alpha$ =2.8 ms / 1.39 ms / 70°, SENSE-rate=2, linear ordering, 10 linear ramp-up pulses and acquisition window=239.8 ms, bandwidth=1845 Hz/pixel. The sliceinterleaved T<sub>2</sub> mapping sequence had the following parameters: 5 short-axis slices, in-plane resolution=2×2 mm<sup>2</sup>, slice thickness=8 mm, slice gap=4 mm, field-of-view=320×320 mm<sup>2</sup>, TR/TE/α=2.8 ms / 1.41 ms / 55°, SENSE-rate=2.5, linear ordering, 10 linear ramp-up pulses and acquisition window=188.7 ms, bandwidth=1785.7 Hz/pixel. Because of limited availability of scan time and the heterogeneity of our patient cohorts, no reproducibility assessments were performed in this cohort.

#### **Data Analysis**

**Reproducibility Study**: All images were transferred to a separate workstation for analysis. T<sub>1</sub> and T<sub>2</sub> mapping of each scan were estimated by voxel-wise curve fitting of the signal after motion correction. In-plane motion between different images of T<sub>1</sub> and T<sub>2</sub> maps were corrected using image registration by a non-rigid image registration algorithm (24).  $T_1$  values were estimated by fitting the recovery curve respectively to a 2-parameter and T<sub>2</sub> values to a 3parameter fitting model (25). The endocardial and epicardial contours of the LV myocardium were manually outlined. The anterior right ventricular (RV) insertion point was marked as a point of reference to generate a 26 segment model of five slices over the entire left ventricle from apex to base. Additionally, a mean  $T_1/T_2$  estimate was generated for each of the five single slices and for the entire ventricle. Artefacts were excluded by manually drawn ROI's and any segment with visual artefacts after segment analysis was excluded from the analysis. A second experienced reader analyzed T<sub>1</sub> and T<sub>2</sub> measurements for one repetition in all subjects - which was compared with results of the corresponding repetition of the first reader - to assess interobserver agreement by using intra-class correlation coefficient (ICC) analysis. Intra-observer agreement was assessed by using intra-class correlation coefficient (ICC) analysis for two corresponding repetitions of each subject analyzed by one reader. Continuous values were presented as mean ± standard deviation (SD). The Shapiro-Wilk test was used to assess for normal distribution. Coefficient of variation analysis (CV) was generated to assess variability between subjects, days and sessions and was visualized by box plots. Significance was considered as p-value less than 0.05. Data analysis was performed with SPSS software (SPSS, Version 17, Inc., Chicago, IL, USA) and Matlab software (The MathWorks Inc., Natick, MA).

**Impacts on Sample Size Calculation:** A linear mixed effects model was used to calculate variance estimations for the effect of volunteers, days, sessions, repetitions and remaining, unspecified factors (i.e. heart rate, etc.). For this model, imaging day is nested within subjects, session is nested within day and repetition is nested within session. This analysis provides individual variances for each individual factor, a pooled variance and pooled standard deviation for each T<sub>1</sub> (slice-interleaved/MOLLI/ShMOLLI) and T<sub>2</sub> (slice-interleaved/single-slice) mapping sequence. The sample size assessment was performed for 90%, 85% and 80% power groups for detection of different changes in T<sub>1</sub> and T<sub>2</sub> values at a type 1 error of 0.05.

**Image Quality Assessment**: Subjective image quality was assessed by two readers (S.B. with 2 years of experience and C.L. with 6 years of experience) using a 26-segment LV model. The LV was divided into five short-axis slices perpendicular to the longitudinal axis of the heart. This leads to five circular sections of the LV-myocardium (basal, three mid-ventricular and apical slices). Only slices with complete circular section of the LV were included. Each of the three most basal sections were divided into six segments of 60° each. Each of the two most apical slices were divided into 4 segments of 90° each. Segments were evaluated independently by two readers. A score of 1 was given if the segment was of acceptable image quality for analysis, defined in this study as having at least a  $5 \times 5$  pixel region of interest (ROI) not affected by artifact. Otherwise, a score of 0 was given to this segment. Inter-observer agreement for both slice-interleaved T<sub>1</sub> and T<sub>2</sub> maps were calculated.

### RESULTS

### **Reproducibility Study**

Figure 2 shows example T<sub>1</sub> mapping images of five short-axis LV slices for the slice-interleaved  $T_1$  sequence with eight repetitions (two days × two sessions × two repetitions). All five slices of the eight repetitions appear with homogeneous quality and without artefacts. Figure 3 is an example of eight repetitions of T<sub>2</sub> mapping of five LV slices for slice-interleaved T<sub>2</sub> sequence. All slices in all repetitions of this subject show homogeneous quality and are without artefacts. Figure 4 shows individual and group global T<sub>1</sub> and T<sub>2</sub> estimates among all 11 healthy subjects at each repetition. Each sequence was repeated twice per session. There were two sessions on each of the two days. The results including standard deviations for both sequences appeared very similar between different repetitions, sessions and days. The global mean T<sub>1</sub> time for sliceinterleaved T<sub>1</sub> was 1063 $\pm$ 22 ms and the global mean T<sub>2</sub> time for slice-interleaved T<sub>2</sub> was 48 $\pm$ 5 ms. Figure 5 shows global T<sub>1</sub> and T<sub>2</sub> estimates for each of the five slices among all repetitions and subjects. The slices within each sequence showed similar global  $T_1$  and  $T_2$  estimates. Segments compromised by severe artefacts were excluded (T<sub>1</sub>: 5.6% and T<sub>2</sub>: 16.3% of excluded segments). Slice-interleaved T<sub>2</sub> showed a higher tendency for motion artefacts than sliceinterleaved T<sub>1</sub>.

The coefficient of variation analysis (CV) between different subjects, days and sessions and for both sequences showed a high reproducibility with global CV's of < 5% between days, sessions and subjects (Figure 6). The CV analysis for individual slices showed a low variability for slice-interleaved T<sub>1</sub> (global CV per slices: slice 1=2.7%; slice 2=2.8%; slice 3=3.2%; slice 4=3.1%; slice 5=3.7%) as well as slice-interleaved T<sub>2</sub> (global CV per slices: slice 1=12.5%; slice 2=11.5%; slice 3=8.8%; slice 4=9.4%; slice 5=8.3%). The most apical and most basal slices

(slice 1 and slice 5) showed similar reproducibility as compared to the three mid-ventricular slices (slice 2-4) in both sequences after exclusion of segments with severe artefacts.

The inter-observer agreement for slice-interleaved  $T_1$  sequence showed an excellent agreement with an ICC of 0.86 (95% confidence interval: 0.13 to 0.97). The inter-observer agreement for slice-interleaved  $T_2$  sequence was very strong with an ICC of 0.75 (95% confidence interval: 0.17 - 0.93). The intra-observer agreement for slice-interleaved  $T_1$  sequence showed an excellent agreement with an ICC of 0.87 (95% confidence interval: 0.35 - 0.97) and the intra-observer agreement for slice-interleaved  $T_2$  sequence was very strong with an ICC of 0.77 (95% confidence interval: 0.12 - 0.94).

### **Impacts on Sample Size Calculation**

**Figure 7** shows the required sample size to detect changes in T<sub>1</sub> measured using STONE, MOLLI and ShMOLLI at 3 different statistical power levels (>90%, >85%, >80%). The required sample size for detection of a specific T<sub>1</sub> difference is smaller for STONE than for MOLLI and ShMOLLI. **Figure 8** shows the required sample size to detect changes in T<sub>2</sub> measured by using slice-interleaved T<sub>2</sub> and single-slice T<sub>2</sub> mapping sequences at three different statistical power levels (>90%, >85%, >80%). For both T<sub>2</sub> sequences, the required sample sizes are very similar for detection of T<sub>2</sub> differences. For detection of T<sub>2</sub> differences of  $\leq$ 10ms, single-slice T<sub>2</sub> requires fewer subjects than slice-interleaved T<sub>2</sub>.

#### **Image Quality Assessment**

Five patients were excluded from analysis due to incorrect positioning of the slices and/or reconstruction error. **Figure 9** shows the subjective image score (averaged between the two readers) for each of the 26 T<sub>1</sub> and T<sub>2</sub> map segment. The lowest scores consistently seen on both T<sub>1</sub> and T<sub>2</sub> maps were at the most basal and most apical slices (slices one and five, respectively) and adjacent to the RV insertion points into the interventricular septum. The mean visual quality scores for the most basal slices were  $0.81\pm0.04$  and  $0.81\pm0.04$  for T<sub>1</sub> and T<sub>2</sub> maps respectively. The mean visual quality scores for the most apical slices were  $0.61\pm0.03$  and  $0.76\pm0.03$  for T<sub>1</sub> and T<sub>2</sub> maps, respectively. The septal wall segments demonstrated the best scores in both T<sub>1</sub> and T<sub>2</sub> maps, with mean visual quality scores of  $0.85\pm0.09$  and  $0.84\pm0.04$  for T<sub>1</sub> and T<sub>2</sub> maps, respectively. Using the Wilcoxon Signed-Rank test, the mean score differences between basal and septal as well as apical and septal segments were significantly different (p<0.001) for T<sub>1</sub> and T<sub>2</sub> maps, respectively.

The inter-observer agreement for visual quality scores of slice-interleaved  $T_1$  maps showed strong agreement with an ICC of 0.74 (95% confidence interval: 0.64 - 0.81). The inter-observer agreement for quality scores of slice-interleaved  $T_2$  sequence was moderate to good with an ICC of 0.60 (95% confidence interval: 0.35 - 0.74).

# DISCUSSION

In this prospective study examining  $T_1$  and  $T_2$  reproducibility with slice-interleaved  $T_1$  and  $T_2$  mapping sequences in healthy subjects and subjective image quality in patients with known or suspected cardiovascular disease, we found that these two free-breathing sequences provide excellent reproducibility. The subjective image quality analysis demonstrated >80% of segments are suitable for quantitative measurements, allowing regional measurements, which are important in certain diseases such as myocarditis and hypertrophic cardiomyopathy.

Our results expand on previous observations regarding the reproducibility of  $T_1$  mapping using different  $T_1$  mapping sequences (18). While SASHA and SAPHIRE have excellent accuracy compared to MOLLI and ShMOLLI, they have lower reproducibility. The STONE sequence has a longer recovery time than MOLLI (18,20), which improves accuracy and precision. Our data also support improved reproducibility of the STONE sequence as the coefficient of variation analysis for STONE was lower than similar studies with MOLLI (26). There are very limited data on reproducibility for myocardial  $T_2$  mapping (27). In a test/retest study involving two separate days, Wassmuth et al. (27) reported a high reproducibility of  $T_2$  mapping with coefficients of variation from 6.6% to 7.6% depending on different imaging orientation.

Over the past several years, there have been significant advances in pulse sequence design for myocardial tissue characterization using  $T_1$  and  $T_2$  mapping. As we embark on the next challenge of utilizing these sequences in larger clinical studies, we should incorporate measurement variability in power calculations for future studies. Our preliminary results demonstrated that the STONE  $T_1$  sequence will lower the sample size needed to achieve a pre-specified power to

Journal of Magnetic Resonance Imaging This article is protected by copyright. All rights reserved. detect changes in  $T_1$ . In addition to advantages of reduced scan-time and free-breathing acquisition compared to other available  $T_1$  mapping sequences, a lower sample size may reduce the overall cost of clinical studies and increase the sensitivity to detect smaller changes in  $T_1$  in longitudinal studies. Our  $T_2$  power calculation shows that similar numbers of patients are needed for the two sequences to detect expected  $T_2$  differences; however, slice-interleaved  $T_2$  will still require a shorter scan time. In our calculation, we did not consider non-diagnostic segments that should be accounted for when planning a clinical study to guarantee sufficient statistical power. Additional studies are warranted to calculate sample sizes in specific population of patients that might contain different degrees of  $T_1$  and  $T_2$  variability.

Our study has several limitations. The reproducibility measurements were only assessed in a small cohort of healthy young adult subjects and the statistical power analysis was mainly based on results in healthy subjects. The reproducibility may be lower in patients with different cardiovascular diseases. For example, in patients with dilated cardiomyopathy, the LV wall is often thin, making it difficult to measure  $T_1$  or  $T_2$  values. In our experience, it is difficult to perform similar test/retest studies on two separate days in patients, therefore we only performed this study in healthy subjects.  $T_1$  and  $T_2$  may be dynamic. In our healthy cohort, we attributed the measurement variability to performance of the imaging sequence rather than to the changes in underlying  $T_1$  or  $T_2$ . Nonetheless, this may not be the case in patients. We did not assess reproducibility of post-contrast  $T_1$  or ECV in our healthy cohort as this would require contrast administration and hematocrit measurement. Additionally, we did not measure hematocrit in our patients, thus we only performed native  $T_1$  sequence.

In conclusion slice-interleaved  $T_1$  and  $T_2$  mapping yield highly reproducible myocardial  $T_1$  and  $T_2$  values, which may have implications for the determination of required sample sizes in larger

clinical studies. Full LV coverage allows for assessment of various myocardial segments, with >80% for T<sub>1</sub> and >83% for T<sub>2</sub> maps of interpretable segments.

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**Figure 1:** Study protocol: Healthy subjects (n=11) underwent MR imaging to assess acquisition reproducibility between days, sessions and repetitions. The study protocol was identical in both days. Each sequence was repeated twice per session. Between two different sessions, subjects were taken out of the scanner, and the 32-channel phased array coil was repositioned before starting the next session.

**Figure 2**: Example T<sub>1</sub> maps acquired with the STONE sequence.

**Figure 3:** Example  $T_2$  maps acquired with the free-breathing slice-interleaved  $T_2$  mapping sequence.

**Figure 4:** Mean  $\pm$  standard deviation (among different subjects) of global myocardial T<sub>1</sub> (A) and T<sub>2</sub> (B) measurements for different repetitions.

**Figure 5:** Mean  $\pm$  standard deviation (among different subjects) of global myocardial T<sub>1</sub> (A) and T<sub>2</sub> (B) estimates for different slices (basal: slice 1, mid-ventricular: slices 2-4, and apical: slice

**Figure 6:** Coefficient of variation (CV) analysis for  $T_1$  (A) and  $T_2$  (B) mapping sequences to assess the variability between different subjects, days and sessions.

**Figure 7:** Assessment of required sample size for detection of corresponding  $T_1$  differences in STONE, MOLLI and ShMOLLI mapping sequences for 3 different power levels (>90%/ >85%/ >80%;  $\alpha$ =0.05).

**Figure 8:** Assessment of required sample size for detection of corresponding  $T_2$  differences in slice-interleaved  $T_2$  and single-slice  $T_2$  mapping sequences for 3 different power levels (>90%/ >85%/ >80%;  $\alpha$ =0.05).

**Figure 9:** A 26-segment polar map of the left ventricle showing the average visual quality scores (0=poor; 1=good) for the two readers using the 26 segment LV model for (a)  $T_1$  mapping and (b)  $T_2$  mapping, respectively.

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Figure 1: Study Protocol: Healthy subjects (n=11) underwent MR imaging to assess acquisition reproducibility between days, sessions and repetitions. The study protocol was identical in both days. Each st, fere. apositio. x 300 DPI) sequence was repeated twice per session. Between two different sessions, subjects were taken out of the scanner, and the 32-channel phased array coil was repositioned before starting the next session. 63x34mm (300 x 300 DPI)

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Figure 2: Example  $T_1$  maps acquired with the STONE sequence. 79x83mm (300 x 300 DPI)













Figure 5: Mean with standard deviation (among different subjects) of global myocardial T<sub>1</sub> (A) and T<sub>2</sub> (B) estimates for different slices (basal: slice 1, mid-ventricular: slice 2-4, and apical: slice 5). 188x259mm (300 x 300 DPI)

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**Figure 7:** Assessment of required sample size for detection of corresponding  $T_1$  differences in sliceinterleaved  $T_1$ , MOLLI and ShMOLLI mapping sequences for groups of 3 different power levels (>90%/ >85%/ >80%; a=0.05). 183x303mm (300 x 300 DPI)







Figure 9: A 26-segment polar map of the left ventricle showing the average visual quality scores (0 poor; 1 good) for the two readers using the 26 segment LV model for (a)  $T_1$  mapping and (b)  $T_2$  mapping, respectively. , 3 75x37mm (300 x 300 DPI)

