

# Fast 3D UTE in vivo $T_1$ and $T_2^*$ mapping of fast relaxing knee tissues at 3 T

Maik Rothe<sup>1,2</sup>  | Selina Riedel<sup>1</sup> | Anne Slawig<sup>1,2</sup>  | Andreas Deistung<sup>1,2</sup>  |  
Klaus Bohndorf<sup>1</sup> | Richard Brill<sup>1</sup> | Walter A. Wohlgemuth<sup>1,2</sup>  | Alexander Gussew<sup>1,2</sup> 

<sup>1</sup>University Clinic and Outpatient Clinic for Radiology, University Hospital Halle (Saale), Halle (Saale), Germany

<sup>2</sup>Halle MR Imaging Core Facility (HMRICF), Halle (Saale), Germany

## Correspondence

Maik Rothe, University Clinic and Outpatient Clinic for Radiology, University Hospital Halle (Saale), Ernst-Grube-Str. 40, D06120 Halle (Saale), Germany.  
Email: [maik.rothe@uk-halle.de](mailto:maik.rothe@uk-halle.de)

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## Abstract

**Purpose:** UTE MR imaging captures quantitative signals in fast-relaxing tissues, enabling anatomical visualization and quantitative assessment of  $T_1$  and  $T_2^*$  relaxation times. However, the clinical application of quantitative UTE MRI is limited by long acquisition times. Therefore, this study introduces a novel UTE-based method for  $T_1$  and  $T_2^*$  mapping, achieving submillimeter resolution in less than 10 min.

**Theory and Methods:** The method employs a dual-echo acquisition for fast  $T_2^*$  mapping, augmented by an additional acquisition with different  $T_1$  weighting. This second scan enables the computation of signal ratios between scans with different  $T_1$ -weighting. These measured signal ratios are then compared to a lookup table containing distinct ratios, corresponding to discrete  $T_1$  values. The approach was validated in phantom solutions mimicking various  $T_1$  and  $T_2^*$  times and applied in vivo to quantify relaxation times across different knee tissue compartments in healthy individuals.

**Results:** The method demonstrated its reliability for  $T_1$  and  $T_2^*$  quantification in rapidly relaxing tissues (1–11 ms). However, it exhibited a tendency to underestimate  $T_2^*$  in skeletal muscle. This limitation arises from the chosen TEs being inadequate to capture slow signal decays. In accordance with the findings of preceding studies, this in vivo study identified three distinct  $T_1$  categories of tissue characterized by short (adipose tissue), moderate (ligaments, tendons, and menisci), and long (skeletal muscle)  $T_1$  values.

**Conclusion:** The presented technique for combined  $T_1$  and  $T_2^*$  mapping enables relaxometry in rapidly relaxing tissues, indicating potential for advanced tissue characterization in clinical settings.

## KEYWORDS

$T_2^*$  mapping, knee joint, quantitative imaging,  $T_1$  mapping, UTE

## 1 | INTRODUCTION

MRI of tendons and ligaments is limited by the rapid transverse relaxation times. Conventional sequences with TEs of several milliseconds yield only weak signals from these tissues. UTE techniques enable imaging of fast-relaxing tissues.<sup>1–3</sup> Quantitative UTE MRI provides access to proton density, relaxation, and magnetization transfer properties, offering insights into bound and free water fractions in collagen<sup>4–6</sup> and enabling detection of physiological and pathological changes due to aging, strain, or injury.<sup>7,8</sup>

The knee joint represents a clinically relevant target for quantitative UTE imaging owing to its complex anatomy and abundance of fast-relaxing tissues. Pathological changes in tendons,<sup>9</sup> ligaments,<sup>10–13</sup> cartilage,<sup>14,15</sup> and menisci<sup>16</sup> can be characterized by  $T_1$  and effective  $T_2^*$  relaxation. Whereas  $T_2^*$  is sensitive to local magnetic field inhomogeneities,  $T_1$  more directly reflects the free water content.<sup>17</sup> Therefore, mapping of both enables more comprehensive tissue assessment. The anterior cruciate ligament (ACL), for instance, is particularly prone to degenerative or traumatic injury.<sup>18–20</sup> Early detection of scarring or partial tears may improve treatment outcomes.

Recent UTE studies performed exponential fitting of multiple echoes for  $T_2^*$  mapping.<sup>21–23</sup>  $T_1$  mapping via inversion recovery (IR) is accurate but time-consuming and suboptimally suited for short  $T_2^*$  tissues due to ineffective inversion. Faster but less precise alternatives include saturation recovery with variable flip angle (FA)<sup>24,25</sup> or variable TR.<sup>22</sup> Clinical adoption of UTE relaxometry is limited by long scan times, especially with 3D radial or spiral encoding. Few studies have simultaneously quantified  $T_1$  and  $T_2^*$  in fast-relaxing knee tissues,<sup>9,22,26</sup> typically achieving 2 mm resolution in about 20 min.

In this study, we propose a time-efficient UTE-based method for combined  $T_1$  and  $T_2^*$  mapping of the knee joint at submillimeter isotropic resolution. The method comprises two UTE scans: a dual-echo acquisition for  $T_2^*$  mapping and a scan with variable  $T_1$  weighting to enable lookup table (LUT)-based  $T_1$  quantification.<sup>27</sup> Validation included phantom experiments and in vivo measurements in healthy volunteers across different knee joint tissues, with comparison to literature values.

## 2 | THEORY

The proposed mapping protocol uses two UTE scans with different parameter settings, complemented by a low-resolution  $B_1^+$  map (Figure 1). The first scan acquires one ultrashort echo ( $S_1$ ) with minimized  $T_1$  contrast. The second scan acquires two echoes ( $S_2$ ,  $S_3$ ) using parameters optimized for  $T_1$  sensitivity and for the

Ernst-angle condition in tissues with a  $T_1$  of approximately 500 ms.

### 2.1 | $T_2^*$ mapping

The spoiled gradient echo signal can be modeled as follows<sup>28</sup>:

$$S = S_0 \sin(\alpha) \frac{1 - e^{-\frac{TR}{T_1}}}{1 - e^{-\frac{TR}{T_1}} \cos(\alpha)} e^{-\frac{TE}{T_2^*}}, \quad (1)$$

where  $S$  is the signal intensity,  $S_0$  is the signal intensity at  $TE = 0$  ms, and  $\alpha$  is the FA.

$T_2^*$  can be calculated by logarithmic ratio of two echoes acquired in the second scan:

$$T_2^* = \frac{TE_3 - TE_2}{\ln(S_2/S_3)}, \quad (2)$$

where  $S_2$  and  $S_3$  are signal intensities at ultrashort ( $TE_2$ ) and moderate ( $TE_3$ ), respectively.  $TE_3$  is selected to satisfy the in-phase condition according to the scanner's operating frequency (123.256 MHz at our 3 T system), assuming a chemical shift of 3.3 ppm between water and fat.<sup>29</sup>

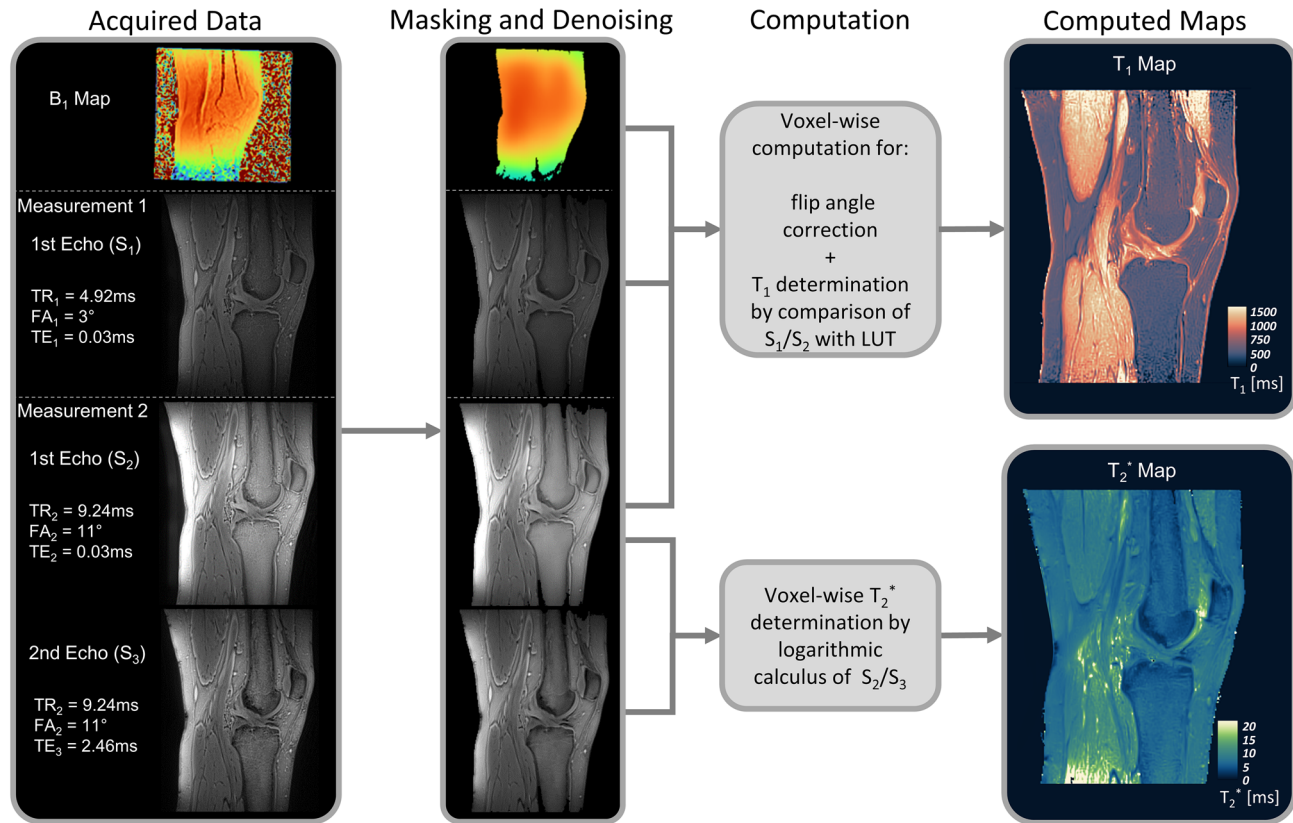
Beyond ensuring in-phase acquisition,  $TE_3$  must also be matched to the expected  $T_2^*$  range. Long  $TE_3$  improves accuracy for long  $T_2^*$  values but reduces it for short  $T_2^*$  and vice versa (Figure S1). Details and visualizations of  $TE_2/TE_3$  optimization are provided in Figure S1. In this study,  $T_2^*$  mapping was evaluated for two different settings of  $TE_3$  (2.46 ms and 4.92 ms) and compared to conventional mono-exponential fitting of three echoes ( $TEs = 0.03/2.46/4.92$  ms).

### 2.2 | $T_1$ mapping

$T_1$  values are derived from two UTE measurements,  $S_1$  and  $S_2$ , acquired with different FA and/or TR, using the signal model described in Equation 1. Because the ratio  $S_1/S_2$  is determined by the underlying  $T_1$  value, we constructed a LUT that maps discrete  $S_1/S_2$  ratios to specific  $T_1$  values within a predefined range<sup>27</sup> (1–4000 ms in 1 ms steps in our study):

$$\frac{S_1}{S_2} = \frac{\sin(\alpha_1) \left(1 - e^{-\frac{TR_1}{T_1}}\right) \left(1 - e^{-\frac{TR_2}{T_1}} \cos(\alpha_2)\right)}{\sin(\alpha_2) \left(1 - e^{-\frac{TR_2}{T_1}}\right) \left(1 - e^{-\frac{TR_1}{T_1}} \cos(\alpha_1)\right)}. \quad (3)$$

Accurate knowledge of the actual FAs is crucial but can be extracted from a low-resolution  $B_1^+$  map.



**FIGURE 1** Overview of the proposed method for T<sub>1</sub> and T<sub>2</sub><sup>\*</sup> mapping, illustrated on a sagittal knee image using the Lipari and Navia color maps.<sup>71</sup> The left column presents the proposed acquisition scheme consisting of two UTE scans and B<sub>1</sub><sup>+</sup> mapping. In the subsequent image preprocessing, the background was removed by masking and the SNR was improved by a denoising algorithm (second column). After these steps, the T<sub>1</sub> and T<sub>2</sub><sup>\*</sup> maps were calculated (third column). The right column shows the corresponding T<sub>2</sub><sup>\*</sup> map (based on logarithmic calculation of S<sub>2</sub> and S<sub>3</sub>) as well as the B<sub>1</sub><sup>+</sup>-corrected T<sub>1</sub> map (based on S<sub>1</sub> and S<sub>2</sub> via LUT). Peripheral T<sub>1</sub> inhomogeneity in muscle (e.g., femur) may reflect transmit-field imperfections at the edge of the FOV. LUT, lookup table; S, signal intensity.

The choice of FA<sub>1</sub>/FA<sub>2</sub> and TR<sub>1</sub>/TR<sub>2</sub> also depends on the expected T<sub>1</sub> range and is constrained by the settings used for T<sub>2</sub><sup>\*</sup> mapping. To reduce scan time, TR<sub>1</sub> and TR<sub>2</sub> are kept as short as possible. FA<sub>2</sub> is selected according to the Ernst angle for tissues with T<sub>1</sub> ≈ 500 ms. This leaves FA<sub>1</sub> as the only remaining parameter to be optimized. It must be small enough to minimize T<sub>1</sub> weighting across a broad T<sub>1</sub> range, while remaining large enough to ensure adequate SNR in the first UTE scan (S<sub>1</sub>). Based on numerical simulations (Figure S1), FA<sub>1</sub> = 3° was chosen as a sufficient compromise.

### 3 | METHODS

The proposed fast T<sub>1</sub> and T<sub>2</sub><sup>\*</sup> mapping approach was evaluated in vitro and applied in vivo in 20 asymptomatic young volunteers (14/6 female/male, 26.6 ± 6.2 years) with no history of knee trauma, pain, functional impairment, or intense sports activity. The study was approved

by the local ethics committee (protocol no.: 2021-056) and conducted in accordance with the Declaration of Helsinki; written informed consent was obtained from all participants.

#### 3.1 | Phantom construction

Two multi-compartment phantoms were constructed to mimic T<sub>1</sub> and T<sub>2</sub><sup>\*</sup> relaxation properties in knee tissues. The T<sub>2</sub><sup>\*</sup> phantom (seven tubes) was designed to simulate T<sub>2</sub><sup>\*</sup> values between 1 ms and 20 ms using 3 wt% carrageenan gels with 0.9 wt% sodium chloride and varying cornstarch concentrations ([80/70/60/50/45/40/33] wt%). Cornstarch effectively shortens T<sub>2</sub><sup>\*</sup> without inducing B<sub>0</sub> field distortions.<sup>30-32</sup> The T<sub>1</sub> phantom (10 tubes) simulated T<sub>1</sub> values between 300 and 1300 ms<sup>33,34</sup> using 3 wt% agarose gels with 0.9 wt% sodium chloride and graded concentrations of gadolinium ([300/180/140/80/60/40/30/20/12/7] μM, gadobutrol).

### 3.2 | Data acquisition

MRI was performed on a clinical 3 T MR scanner (Magnetom Vida, Siemens Healthineers, Erlangen, Germany) using an 18-channel transmit/receive knee coil. In vivo scans were acquired with volunteers in supine position, with knees fixated and angulated by approximately 15°.

The imaging protocol included two UTE scans with 3D stack-of-spirals readout (prototype spoiled gradient echo UTE sequence<sup>35,36</sup>) and a low-resolution  $B_1^+$  map (Figure 1). Scans were performed in sagittal orientation with 0.8 mm isotropic resolution (20  $\mu$ s block-pulse excitation, FOV: 198  $\times$  198 mm<sup>2</sup>, matrix: 256  $\times$  256, 172 slices, slice thickness: 0.8 mm). Each slice was encoded with 512 spiral readouts (1160  $\mu$ s readout, 682 samples, spectral bandwidth: 588 kHz, pixel bandwidth: 2298 Hz/pixel, TE range: 30–660  $\mu$ s from k-space center to periphery).

- Measurement 1 (3:18 min): TE: 0.03 ms, TR<sub>1</sub>: 4.92 ms, FA<sub>1</sub>: 3°
- Measurement 2 (5:56 min): TE: 0.03/2.46/4.92 ms, TR<sub>2</sub>: 9.24 ms, FA<sub>2</sub>: 11°

$B_1^+$  mapping was performed using a low-resolution 2D multi-slice turboFLASH sequence<sup>37,38</sup> (TR/TE: 29660/2.56 ms, FA<sub>1</sub>/FA<sub>2</sub>: 8/80°, FOV: 200  $\times$  200 mm,<sup>2</sup> acquisition matrix: 96  $\times$  96, slice thickness: 2 mm). To verify suitability for FA correction, additional experiments in a large cylindrical phantom were performed using the employed  $B_1^+$  mapping sequence (Figure S2).

As a reference for  $T_2^*$  quantification across a broad  $T_2^*$  range, the  $T_2^*$  phantom was scanned using an echo-train shifted multi-echo (ETsME) approach. Here, 22 measurements were repeated with the first echo shifted from TE<sub>1</sub>: 30–1500  $\mu$ s, while keeping the subsequent echoes constant (TE<sub>2-5</sub>: 4.92/7.38/9.84/12.3 ms, TR: 13 ms; FA: 12°). Reference  $T_1$  values of the  $T_1$  phantom were obtained via inversion-prepared UTE scans (TE/TR: 0.03/4.4 ms, FA: 6°, 22 TIs: 30–8000 ms) and fitted using three free parameters.

### 3.3 | Data preprocessing

The low-resolution  $B_1^+$  map was resampled and aligned to the high-resolution UTE images using the FreeSurfer package (version 7.4.0) (<http://surfer.nmr.mgh.harvard.edu><sup>39</sup>). Subsequent processing was performed with custom Python scripts using standard libraries.<sup>40</sup> To mitigate FA uncertainty in fast-relaxing tissues, the  $B_1^+$  map was polynomially smoothed and used to compute voxelwise FA correction factors. UTE images were denoised using an

adaptive nonlocal means filtering algorithm<sup>41</sup> and affinely coregistered. Finally, subtraction images ( $S_2$ – $S_3$ ) were generated to highlight fast-relaxing tissue structures.

$T_2^*$  reference values in phantoms were determined via monoexponential fitting of the ETsME decay.  $T_2^*$  mapping in both phantom and in vivo data was performed voxelwise using three different approaches:

- 3TE: Three-point mono-exponential fit using TEs: 0.03/2.46/4.92 ms
- 2TE<sub>2,46</sub>: Dual-echo computation (Equation (2)) using TE<sub>2</sub>/TE<sub>3</sub>: 0.03/2.46 ms
- 2TE<sub>4,92</sub>: Dual-echo computation (Equation (2)) using TE<sub>2</sub>/TE<sub>3</sub>: 0.03/4.92 ms

$T_1$  mapping was performed via a LUT generated from simulated  $S_1/S_2$  signal ratios corresponding to  $T_1$  values between 1 ms and 4000 ms (Figure S1D). To account for  $B_1^+$  inhomogeneity, separate LUTs were generated for FAs from 1° to 180° (1° steps). Each voxel's local FA was derived from the  $B_1^+$  map and used to select the corresponding LUT. Reference  $T_1$  values in the phantom were determined by monoexponential fitting of the IR series.

### 3.4 | Volumes of interest

Volumes of interest (VOIs) were manually defined using 3D Slicer (<https://www.slicer.org><sup>42</sup>) based on subtraction images ( $S_2$ – $S_3$ ) and transferred to the parameter maps to extract VOI specific mean values of relaxation parameters.

Two circular VOIs per phantom tube were placed on matching slices (~3 pixels from the tube wall) and interpolated across slices to form 3D cylindrical VOIs.

In vivo, VOIs comprising at least 100 voxels were manually segmented for nine knee tissues by consensus of four raters (S.R., M.R., K.B., A.G.) based on sagittal, coronal, and axial views (Figure S3). Tissues included the ACL, posterior cruciate ligament (PCL), patellar tendon (PT), quadriceps tendon (QT), posterior horn of lateral meniscus (LM), bone marrow (BM), infrapatellar fat pad (IFP), subcutaneous adipose tissue (SAT), and skeletal muscle (SM). For the meniscus, tendons, and ligaments, only hyperintense voxels in subtraction images were selected to emphasize fast-relaxing components; vessels and fascia were excluded whenever possible.

### 3.5 | Statistical analysis

Agreement between the proposed and reference methods for  $T_2^*$  and  $T_1$  quantification in phantoms was evaluated using Bland–Altman analysis.  $T_2^*$  values derived from the



3TE, 2TE<sub>2.46</sub>, and 2TE<sub>4.92</sub> approaches were assessed in relation to those from ETsME series. LUT-based  $T_1$  values were evaluated against values obtained from the IR method. Limits of agreement were defined as the mean difference  $\pm 1.96$  SD.

One-way analysis of variance with Tukey's post hoc test was used to evaluate tissue-specific differences in in vivo  $T_1$  and  $T_2^*$  values.  $T_2^*$  assessment was primarily based on 2TE<sub>2.46</sub> data. A separate analysis of variance tested for method-related differences in  $T_2^*$  values across tissues.

All analyses were performed using GraphPad Prism (version 10.0.0 for Windows, GraphPad Software, Boston,

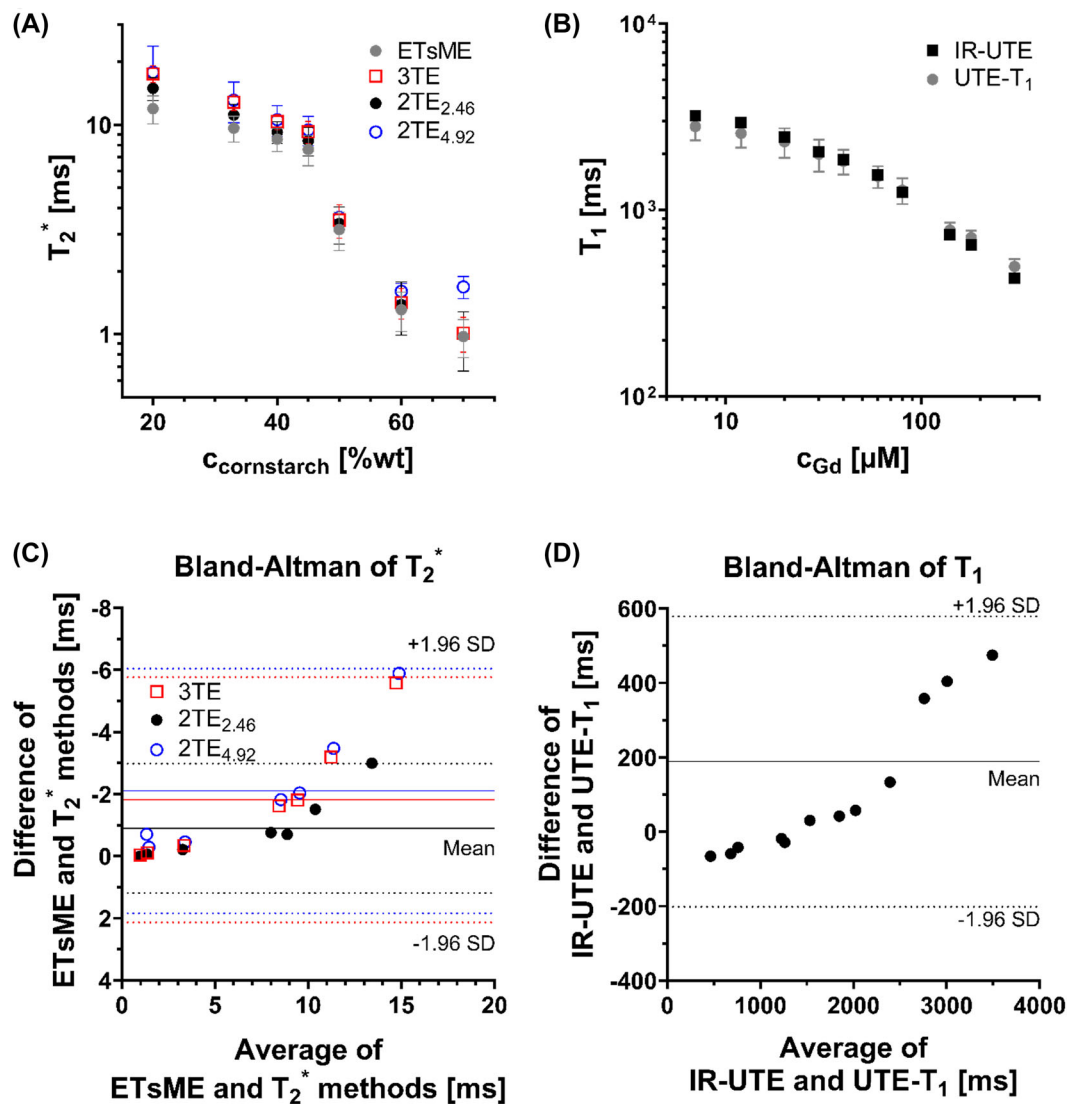
MA, [www.graphpad.com](http://www.graphpad.com)). A  $p$ -value  $< 0.05$  was considered statistically significant.

## 4 | RESULTS

### 4.1 | Phantom experiments

As expected,  $T_1$  and  $T_2^*$  values decreased in the phantoms with increasing gadolinium and cornstarch concentrations, respectively (Figure 2).

$T_2^*$  values derived from all three UTE-based methods closely matched ETsME references, showing mean  $\pm$  SD



**FIGURE 2** Quantitative mapping in phantom experiments. Mean  $T_2^*$  (A) and  $T_1$  (B) values are plotted for different concentrations of cornstarch and gadolinium, respectively, for the proposed fast UTE mapping technique and the corresponding reference methods (IR-UTE for  $T_1$  and ETsME UTE for  $T_2^*$ ). The corresponding Bland-Altman plots are shown in the bottom row for  $T_2^*$  (C) and  $T_1$  (D). The differences in (C) and (D) were calculated as reference method - mapping method. The solid and dotted lines indicate the mean and  $\pm 1.96$  SD of the differences between the reference method and the UTE-based method, respectively (red: 3TE, black: 2TE<sub>2.46</sub>, blue: 2TE<sub>4.92</sub>). ETsME, echo-train shifted multi-echo.

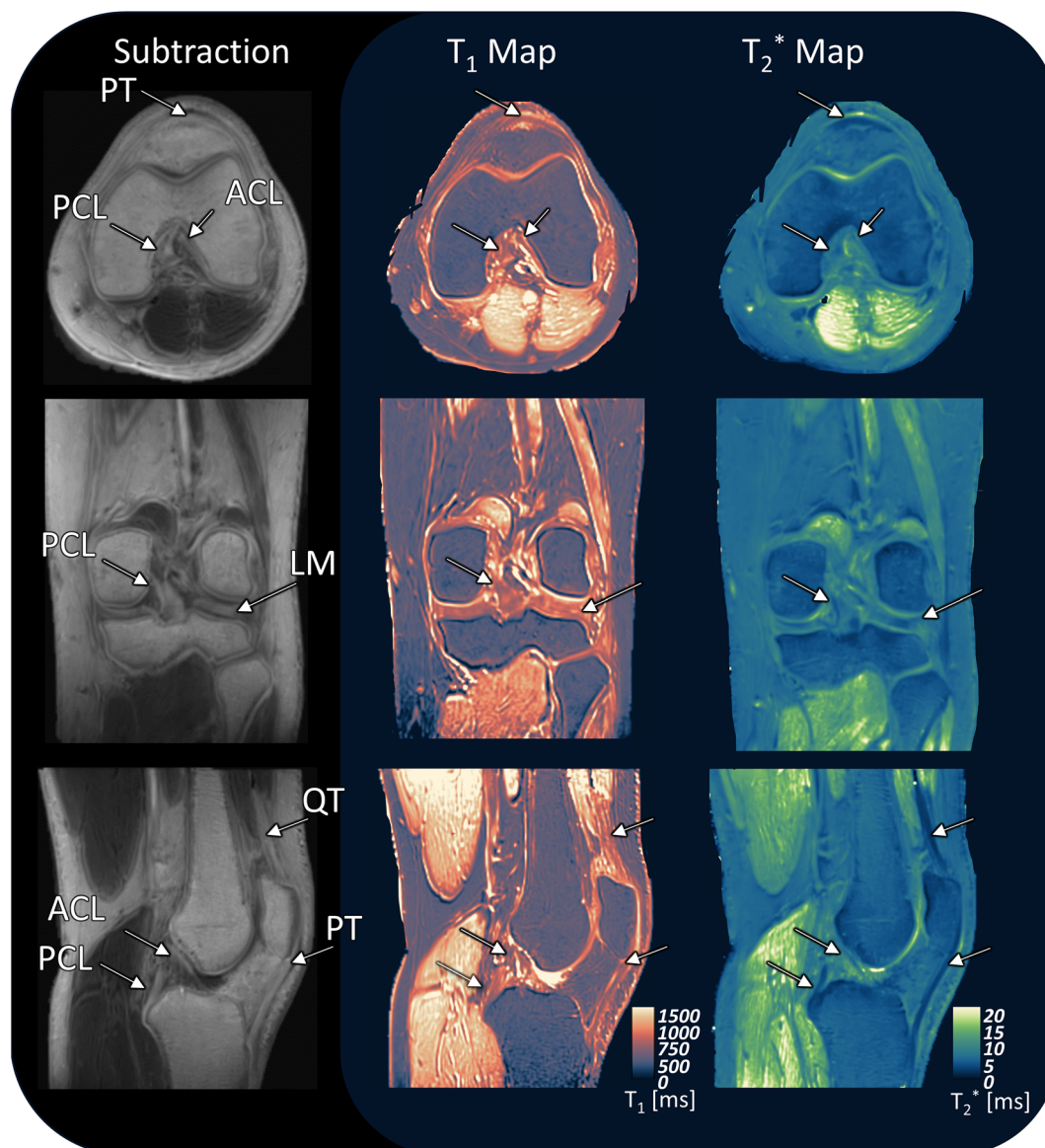
differences of  $16\% \pm 10\%$  (3TE),  $9\% \pm 7\%$  ( $2TE_{2.46}$ ), and  $24\% \pm 11\%$  ( $2TE_{4.92}$ ) (Figure 2A). Bland-Altman analysis (Figure 2C) demonstrated good agreement up to  $T_2^* = 11$  ms; beyond this,  $2TE_{4.92}$ , and 3 TE increasingly overestimated  $T_2^*$ . Additionally,  $2TE_{4.92}$  systematically overestimated values below 3 ms. No reliable values were obtained for  $T_2^* < 1$  ms.

LUT-based  $T_1$  values closely matched IR-UTE references (Figure 2B), with strong agreement up to  $T_1 = 2500$  ms (mean difference:  $2\% \pm 7\%$ , maximum

deviation: 9%). For  $T_1 > 2500$  ms, LUT estimates exhibited reduced accuracy with a discrepancy of  $13\% \pm 1\%$  (Figure 2D).

## 4.2 | In vivo experiments

Representative UTE subtraction images and corresponding  $T_1$  and  $T_2^*$  maps are shown in Figure 3, illustrating signal behavior and regional contrast in a healthy knee.



**FIGURE 3** Typical images of the proposed UTE mapping method in three orientations from one volunteer. The subtraction images ( $S_2 - S_3$ ) are shown in the left column, whereas the generated  $T_1$  maps (displayed in Lipari colormap<sup>71</sup>) and  $T_2^*$  maps from the dual-echo method ( $2TE_{2.46}$ , displayed in Navia colormap<sup>71</sup>) are shown in the middle and right columns, respectively. Arrows mark the posterior horn of the LM, ACL, PCL, QT, and PT. ACL, anterior cruciate ligament; LM, lateral meniscus; PCL, posterior cruciate ligament; PT, patellar tendon; S, signal intensity; QT, quadriceps tendon.

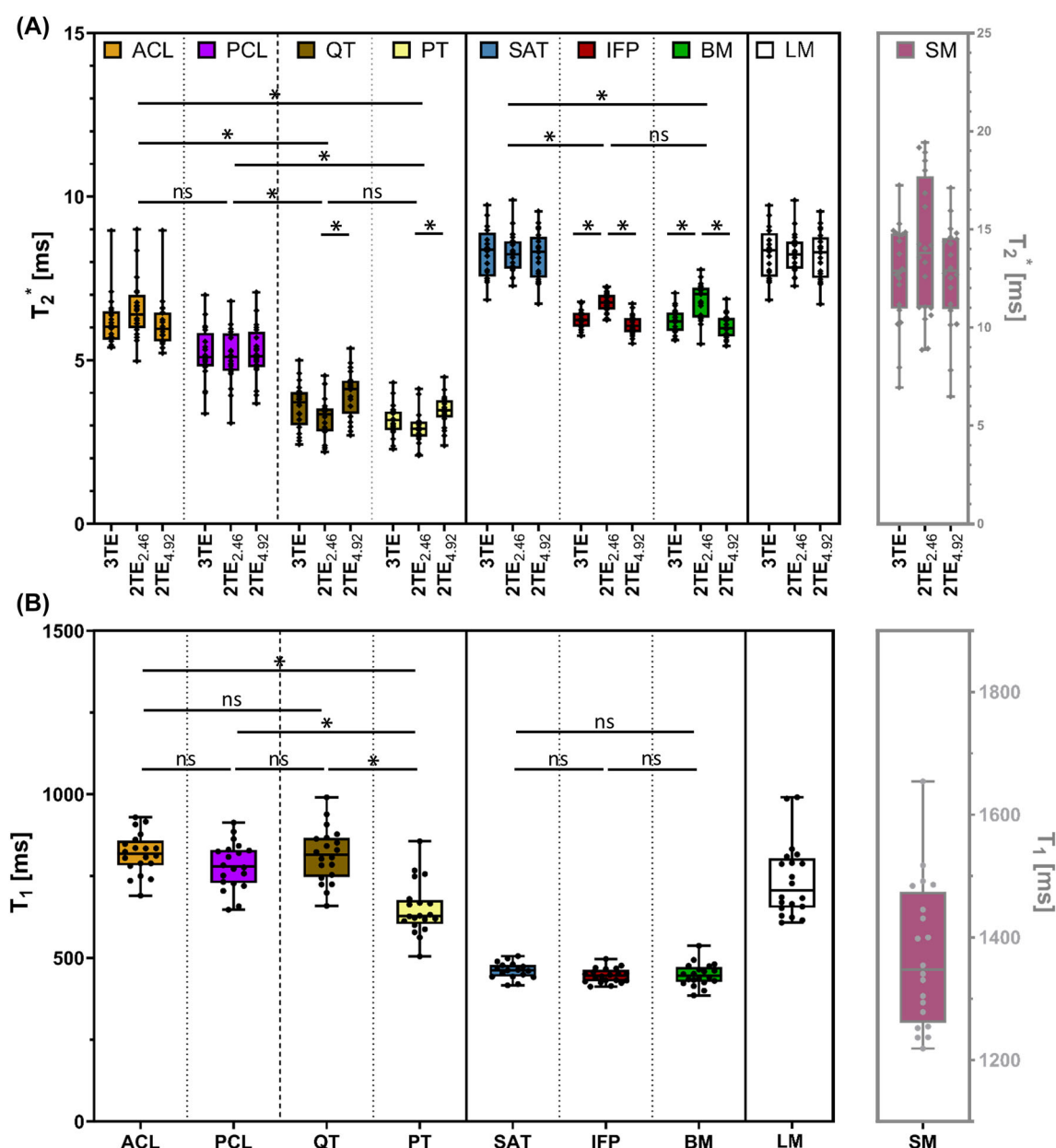
Figure 4 and Table 1 summarize the quantitative  $T_1$  and  $T_2^*$  values across nine examined tissue compartments, alongside literature values.

Tendons showed lower  $T_2^*$  values (3–4 ms) than ligaments (5–6 ms), with no significant differences between PT and QT or between ACL and PCL. LM  $T_2^*$  was about 8 ms. Adipose tissues—including SAT, BM, and IFP—had  $T_2^*$  values of 6–8 ms, with significantly higher values in SAT. Interindividual variability (coefficient of variation, CV) was lower in LM and adipose tissues (CV <9%) than in tendons and ligaments (CV: 10%–25%). SM  $T_2^*$  was about

half of the literature values (Table 1), with considerable interindividual variability (CV: 10%–25%).

As demonstrated in the  $T_2^*$  phantom experiments, all three mapping methods yielded consistent in vivo values (Figure 4A). However, the 2TE<sub>2,46</sub> method produced slightly elevated  $T_2^*$  values in the IFP and BM, and lower values in PT and QT.

All measured in vivo  $T_1$  values were below 2500 ms (Figure 4B), which is within the range validated in the  $T_1$  phantom. Tissues clustered into three characteristic  $T_1$  groups: (i) adipose tissue with short  $T_1$  (400 ms–500 ms,



**FIGURE 4** Boxplots of (A)  $T_2^*$  values from three different mapping methods and (B)  $T_1$  values across nine knee tissue types in 20 healthy volunteers. Each circle represents an individual measurement. Statistical significance between tissue types was assessed using one-way ANOVA with Tukey's post hoc test (\* $p < 0.05$ ). ANOVA, analysis of variance; BM, bone marrow; IFP, infrapatellar fat pad; ns, not significant; SAT, subcutaneous adipose tissue; SM, skeletal muscle.

**TABLE 1**  $T_1$  and  $T_2^*$  values for different tissues of the 20 knee joints of healthy volunteers obtained using the proposed imaging protocol compared to literature values.

Tissue	$T_1$ [s]	$T_1$ literature [s]	$T_2^*$ [ms]			$T_2^*$ literature [ms]
			3TE	2TE <sub>2.46</sub>	2TE <sub>4.92</sub>	
Skeletal muscle	1370 ± 120	1060–1420 <sup>33,49,61,68</sup>	12.8 ± 2.6	14.0 ± 3.5	12.7 ± 2.7	24–32 <sup>56</sup>
Bone marrow	450 ± 35	340–380 <sup>24,33</sup>	6.2 ± 0.4	6.8 ± 0.6	6.0 ± 0.4	2–10.3 <sup>22,51,52,57</sup>
Infrapatellar fat pad	450 ± 25	370–400 <sup>33,68a</sup>	6.3 ± 0.3	6.8 ± 0.3	6.1 ± 0.4	5–12.5 <sup>22,53,54a</sup>
Subcutaneous adipose tissue	460 ± 25	370–400 <sup>33,68</sup>	8.3 ± 0.8	8.3 ± 0.6	8.2 ± 0.7	5–12.5 <sup>22,53,54</sup>
Patella tendon	650 ± 85	505–660 <sup>22,49,55,61</sup>	3.2 ± 0.5	3.0 ± 0.5	3.5 ± 0.5	1.6–6.4 <sup>22,58,60,62</sup>
Quadriceps tendon	810 ± 85	700–800 <sup>22,49</sup>	3.6 ± 0.7	3.2 ± 0.6	4.0 ± 0.7	1.4 <sup>22</sup>
Posterior cruciate ligament	780 ± 75	710–840 <sup>49,55,61</sup>	5.2 ± 0.8	5.1 ± 0.8	5.2 ± 0.8	8.3–8.8 <sup>21,55,60</sup>
Anterior cruciate ligament	820 ± 65	740–925 <sup>49,55,61</sup>	6.0 ± 0.5	6.3 ± 0.6	6.0 ± 0.5	9.1–16.3 <sup>11,55,60</sup>
Lateral Meniscus	740 ± 115	600–970 <sup>49,55,61,70,73</sup>	8.3 ± 0.8	8.3 ± 0.7	8.2 ± 0.8	5–10 <sup>3,8,16,60,73</sup>

<sup>a</sup>Assuming same values for infrapatellar fat pad and subcutaneous adipose tissue.

CV: 5%–8%); (ii) tendons, ligaments, and menisci with moderate  $T_1$  (600 ms–900 ms, CV: 8%–16%); and (iii) SM with a long  $T_1$  (>1200 ms, CV: 9%). Within the second group, PT showed significantly lower  $T_1$  than the ligaments ( $p < 0.0001$ ).

## 5 | DISCUSSION

We present a framework for fast quantitative UTE MR imaging that enables combined  $T_1$  and  $T_2^*$  mapping in fast-relaxing musculoskeletal tissues. All required data were acquired in under 10 min at an isotropic resolution of 0.8 mm<sup>3</sup>. The approach can be further accelerated using techniques such as k-space undersampling<sup>43,44</sup> or artificial intelligence-based superresolution,<sup>45,46</sup> thereby further enhancing its clinical applicability. This combination of multiparametric mapping, high resolution, and short scan time distinguishes the method from previous approaches.

Typical  $T_2^*$  mapping requires ≥3 echoes and scan times of 9–20 min,<sup>8,21,22,44,47</sup> often at relatively low (2 mm) or anisotropic resolution, limiting accuracy in small, angled structures such as tendons or menisci due to partial volume effects. Conventional  $T_1$  mapping using multiple TIs,<sup>48</sup> variable FAs,<sup>22,49</sup> or TRs<sup>22</sup> is similarly time-consuming (5–20 min) and spatially limited (~2 mm).

Despite these constraints, multi-parametric approaches—for example, multi-exponential  $T_2^*$  fits of multiple echoes—can enable more precise differentiation of tissues with distinct relaxation properties but are often impractical for routine clinical use. In contrast, our

method specifically targets relevant relaxation ranges via optimized acquisition parameter combinations. The simulations and phantom experiments in this study were designed to define the sensitivity ranges and validate accuracy across different acquisition settings, such as varying TE<sub>3</sub> in dual-echo  $T_2^*$  mapping (2TE<sub>2.46</sub> vs. 2TE<sub>4.92</sub>) or adjusting FA<sub>1</sub> in the supplemental UTE scan for  $T_1$  estimation.

LUT-based  $T_1$  mapping produced accurate values up to 2500 ms, closely matching the IR-UTE reference. This confirms that simplified modeling, when paired with tailored acquisition parameters optimized for each relaxation regime, yields robust  $T_1$  estimates in fast- as well as moderate-to-long relaxing musculoskeletal tissues.

Phantom experiments demonstrated that selecting 2.46 ms as the moderate TE in the dual-echo approach enables more accurate  $T_2^*$  values in fast-relaxing tissues (≤11 ms), with close agreement to reference ETsME fits. Using 4.92 ms led to systematic overestimation for very short  $T_2^*$  values (<3 ms), likely due to excessive signal decay exceeding the measurable dynamic range. Neither method was suitable for extremely short  $T_2^*$  (e.g., cortical bone <1 ms<sup>50</sup>) because signal decay was almost complete at TE = 2.46 ms.

### 5.1 | In vivo experiments

As in the phantom experiments, we evaluated 2TE<sub>2.46</sub>, 2TE<sub>4.92</sub>, and 3 TE  $T_2^*$  mapping in knee tissues. For meniscus and adipose tissues,  $T_2^*$  values aligned well with literature references.<sup>3,8,56,57,16,21,22,51–55</sup> The choice of the moderate echo significantly affected  $T_2^*$  in tendons



( $p < 0.005$ ):  $2TE_{2,46}$  yielded values closest to literature, whereas  $2TE_{4,92}$  and  $3TE$  increasingly overestimated  $T_2^*$ , likely due to advanced signal decay at  $TE_3 = 4.92$  ms. Even  $2TE_{2,46}$  overestimated  $T_2^*$  in the patellar tendon compared to literature values below 3 ms.<sup>22,58</sup> This likely reflects the mono-exponential model's inability to resolve coexisting relaxation components—specifically collagen-bound and free water—thereby masking the fast-relaxing component of the signal in collagen-rich tissue.<sup>59</sup> Whereas multi-exponential models can separate such compartments,<sup>58</sup> they require longer scan times, limiting their clinical practicality.

Other studies reported higher  $T_2^*$  values for ACL and PCL than observed in our cohort,<sup>11,21,55,60</sup> likely due to different segmentation strategies: We specifically targeted fast-relaxing compartments that showed high signal in UTE subtraction images (Figure 2), whereas others analyzed entire ligament volumes, including both collagen-rich and slower-relaxing regions.<sup>49,55,60,61</sup>

Consistent with previous studies,<sup>11,21,22,55,60,62</sup> our analysis confirmed lower  $T_2^*$  values in patellar and quadriceps tendons compared to the ACL and PCL, likely due to their higher collagen content ( $>95\%$  vs.  $<90\%$ ).<sup>59</sup> Higher  $T_2^*$  in ligaments may also reflect their greater angulation relative to  $B_0$ , consistent with the “magic angle” effect.<sup>63–67</sup> In our cohort, tendon and ligament angles ranged between  $20^\circ$  and  $35^\circ$ , which—based on data from Wu et al.<sup>66</sup> on the Achilles tendon—can increase  $T_2^*$  by several milliseconds.

All applied  $T_2^*$  mapping methods underestimated skeletal muscle values due to limited sensitivity to slow-relaxing tissues,<sup>56</sup> necessitating longer TEs for accurate assessment.<sup>8,21,47</sup>

$T_2^*$  values are anatomy- and condition-dependent and poorly differentiate collagen-rich from fatty tissues (Figure 4). In contrast,  $T_1$ —driven by molecular mobility—better reflects tissue properties such as water and macromolecular content. Consistent with previous studies, our results delineate three tissue-specific  $T_1$  categories (Figure 4): short  $T_1$  in adipose tissue (high lipid content)<sup>24,33,68</sup>; moderate  $T_1$  in collagen-rich ligaments, tendons, and menisci<sup>49,55,61,69,70</sup>; and prolonged  $T_1$  in skeletal muscle (elevated free water content).<sup>33,49,61,68</sup> Notably, patellar and quadriceps tendons also differed in  $T_1$ , consistent with Krämer et al.,<sup>22</sup> presumably due to differences in free water and collagen content.

Adipose tissues showed higher  $T_1$  values than literature references (Table 1), likely due to weaker  $T_1$  weighting in  $S_1$  for compounds with  $T_1 < 500$  ms (Figure S1C), consistent with phantom data (Figure 2B/D). Lowering  $FA_1$  in the  $S_1$  scan could enhance accuracy but reduce SNR, requiring systematic assessment of this tradeoff.

## 5.2 | Limitations

A limitation of our  $T_1$  mapping approach is the vendor-specific 2D  $B_1^+$  map,<sup>38</sup> which introduces FA uncertainty in fast-relaxing tissues. Polynomial smoothing was applied to reduce these effects, though small residual errors still may persist. Heterogeneous  $T_1$  distributions in the FOV periphery (e.g., in bone marrow or muscles; Figure 3) likely reflect incomplete FA correction. This inaccuracy was also evident in phantom measurements, where LUT-based  $T_1$  estimates deviated by  $\sim 10\%$  from reference IR values near the FOV edge (Figure S2). Therefore, future work should explore  $B_1^+$  mapping approaches more closely aligned with UTE protocols.<sup>24,49</sup>

Chemical shift artifacts ( $\sim 1$  pixel) at water-fat boundaries—particularly at muscle-fat interfaces (Figure 1) and in thin structures such as tendons, ligaments, and cortical bone—can affect  $T_1$  and  $T_2^*$  mapping by introducing heterogeneous signal contributions in boundary voxels due to partial volume and off-resonance effects. Slightly elevated  $T_2^*$  values in tissues such as the patellar tendon may partly reflect lipid contamination. Fat suppression (e.g., Dixon, spectral saturation) could mitigate this but was not applied due to: (1) reduced efficiency from  $B_0$  inhomogeneities, (2) prolonged scan time, and (3) limited spectral separation at short readouts ( $\sim 1.16$  ms). Future work should therefore develop and evaluate fat suppression techniques optimized for UTE relaxometry.

In stack-of-spirals UTE, the effective TE varies across k-space, from  $30\ \mu\text{s}$  centrally to  $>600\ \mu\text{s}$  peripherally, due to the increasing duration of slice-encoding gradients. Whereas most signal is acquired at short TE, this spread can bias  $T_2^*$  estimates for ultrashort components ( $<2$  ms), thereby reducing contrast or blurring fast-decaying signals. Thus, radial 3D UTE trajectories, which maintain uniform TE across k-space, may offer a more robust alternative for future studies.

## 6 | CONCLUSION

We present a fast framework for submillimeter  $T_1$  and  $T_2^*$  mapping of fast-relaxing tissues. Whole-knee coverage was achieved in under 10 min, providing accurate estimates for  $T_1$  up to 2500 ms and  $T_2^*$  from 1 ms to 11 ms.  $T_1$  mapping effectively differentiated adipose, collagen-rich, and water-rich tissues. Combined  $T_1/T_2^*$  mapping improves overall tissue characterization and may enhance the assessment of small structures such as ligaments and tendons in future studies.

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## ORCID

Maik Rothe  <https://orcid.org/0009-0001-8533-7520>

Anne Slawig  <https://orcid.org/0000-0003-1412-244X>

Andreas Deistung  <https://orcid.org/0000-0002-2427-1302>

Walter A. Wohlgemuth  <https://orcid.org/0000-0002-8076-6000>

Alexander Gussew  <https://orcid.org/0000-0003-2300-4883>

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## SUPPORTING INFORMATION

Additional supporting information may be found in the online version of the article at the publisher's website.

**Figure S1.** Simulation-based optimization of echo times and flip angles for dual-echo  $T_2^*$  and LUT-based  $T_1$  mapping in UTE MRI. (A) Simulated gradient-echo

signal decay curves for different  $T_2^*$  values. Vertical lines mark the ultrashort (0.03 ms) and the first two in-phase echoes at 3 T (2.46 ms, 4.92 ms), assuming a 3.3 ppm water-fat chemical shift at 123.256 MHz. In this study,  $TE_2 = 0.03$  ms and  $TE_3 = 2.46$  ms or 4.92 ms were used. (B) Ratio  $S_2/S_3$  as a function of  $T_2^*$  for different  $TE_3$  values ( $TE_2$  fixed at 0.03 ms). The shaded area indicates the optimal sensitivity range:  $S_3$  decayed by  $\geq 25\%$  for better discrimination of longer  $T_2^*$ , but retained  $>5\%$  of  $S_2$  to reduce noise sensitivity. (C) Simulated gradient-echo signals for ultrashort  $TE = 0.03$  ms with varying  $T_1$  and FA.  $S_2$  ( $TR = 9.24$  ms,  $FA_2 = 11^\circ$ ) corresponds to the  $T_2^*$  mapping scan;  $S_1$  ( $TR = 4.92$  ms,  $FA_1 = 1-6^\circ$ ) corresponds to the first UTE scan with minimized  $T_1$  contrast. (D) Ratio  $S_1/S_2$  versus  $T_1$  for different  $FA_1$  values. The dashed line marks  $FA_1 = 3^\circ$ , chosen as a compromise between minimal  $T_1$  weighting across a wide  $T_1$  range and adequate SNR in  $S_1$ . Shaded limits indicate where  $S_1$ 's  $T_1$  weighting falls below the noise level of  $S_2$  or where  $S_1$  and  $S_2$  are equal within noise.

**Figure S2.** Effects of  $B_1^+$ -correction on  $T_1$ -mapping. Top left: Direct comparison of  $T_1$ -mapping without  $B_1^+$ -correction, with  $B_1^+$ -correction using the dual-angle (DA) approach (as used in this study), with  $B_1^+$ -correction using the Actual Flip angle Imaging (AFI)  $B_1^+$ -mapping approach, and with a gold-standard IR-UTE-based  $T_1$ -mapping experiment. Top right: Three-plane view of the used phantom with a known  $T_1$  of 100 ms<sup>72</sup> (top: axial view; middle: coronal view; bottom: sagittal view). The cylindrical phantom has a height of 20 cm and a diameter of 13 cm and consists of: 3.75 g  $NiSO_4$  and 5 g NaCl per 1000 g  $H_2O$ . Six different cubic volumes (15 mm  $\times$  15 mm  $\times$  15 mm) were positioned in the isocenter along the direction of the main magnetic field. Bottom first row:  $B_1^+$  maps obtained with the AFI (left) and DA (right) methods, displayed in % of nominal FA. The yellow box marks the cropped phantom region used for all maps and includes an overlay of a representative in vivo knee image (50% transparency) to illustrate correspondence with the in vivo field of view. Bottom second row: Corresponding  $T_1$  maps are shown without  $B_1^+$  correction, with DA correction, and with AFI correction. Uncorrected maps exhibit pronounced spatial inhomogeneity, which is reduced by either correction approach.

**Figure S3.** Visualization of manually segmented regions for the patellar tendon, quadriceps tendon, anterior cruciate ligament (ACL), posterior cruciate ligament (PCL), infrapatellar fat pad, subcutaneous adipose tissue, bone marrow, skeletal muscle and the posterior horn of the lateral meniscus. Segments are overlaid on UTE subtraction



images ( $S_1 - S_2$ ) of a knee in axial and three sagittal views. All three planes (axial, coronal, and sagittal) were used for tissue segmentation, as exemplified for the ACL in the three images in the center column of the bottom row. Numbers indicate the planes specified by the dashed lines in the upper left corner.

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