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# **T1ρ Hip Cartilage Mapping in Assessing Patients With Cam Morphology: How Can We Optimize the Regions of Interest?**

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#### Abstract

*Background* T1 $\rho$  MRI has been shown feasible to detect the biochemical status of hip cartilage, but various regionof-interest strategies have been used, compromising the reproducibility and comparability between different institutions and studies.

*Questions/purposes* The purposes of this study were (1) to determine representative regions of interest (ROIs) for cartilage T1 $\rho$  mapping in hips with a cam deformity; and (2) to assess intra- and interobserver reliability for cartilage T1 $\rho$  mapping in hips with a cam deformity.

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K. S. Rakhra, G. Melkus Department of Radiology, University of Ottawa, Ottawa, Canada Methods The local ethics committee approved this prospective study with written informed consent obtained. Between 2010 and 2013, in 54 hips (54 patients), T1p 1.5-T MRI was performed. Thirty-eight hips (38 patients; 89% male) with an average age of  $35 \pm 7.5$  years (range, 23–51 tears) were diagnosed with a cam deformity; 16 hips (16 patients; 87% male) with an average age of  $34 \pm 7$  years (range, 23-47 years) were included in the control group. Of the 38 patients with a cam deformity, 20 were pain-free and 18 symptomatic patients underwent surgery after 6 months of failed nonsurgical management of antiinflammatories and physical therapy. Exclusion criteria were radiologic sings of osteoarthritis with Tönnis Grade 2 or higher as well as previous hip surgery. Three region-of-interest (ROI) selections were analyzed: Method 1: as a whole; Method 2: as 36 to 54 small ROIs (sections of 30° in the sagittal plane and 3 mm in the transverse plane); Method 3a: as six ROIs (sections of 90° in the sagittal plane and one-third of the acetabular depth in the transverse plane: the anterosuperior and posterosuperior quadrants, divided into lateral, intermediate, and medial thirds); and Method 3b: as the ratio (anterosuperior over posterosuperior quadrant). ROIs in Method 3 represent the region of macroscopic cartilage damage, described in intraoperative findings. To asses interobserver reliability, 10 patients were analyzed by two observers (HA, GM). For intraobserver reliability, 20 hip MRIs were analyzed twice by one observer (HA). To assess interscan reliability, three patients underwent two scans within a time period of 2 weeks and were analyzed twice by one observer (HA). T1p values were compared using Student's t test. Interclass correlation coefficient (ICC) and root mean square coefficient of variation (RMS-CV) were used to analyze intraobserver, interobserver, and interscan reliability.

*Results* Patients with a cam deformity showed increased T1p values in the whole hip cartilage (mean:  $34.0 \pm 3.8$  ms versus  $31.4 \pm 3.0$  ms; mean difference: 2.5; 95% confidence interval [CI], 4.7-0.4; p = 0.019; Method 1), mainly anterolateral (2), in the lateral and medial thirds of the anterosuperior quadrant (mean:  $32.3 \pm 4.9$  ms versus 29.4  $\pm$  4.1 ms; mean difference: 3.0; 95% CI, 5.8–0.2; p = 0.039 and mean  $36.5 \pm 5.6$  ms versus  $32.6 \pm 3.8$  ms; mean difference: 3.8; 95% CI, 6.9-0.8; p = 0.014), and in the medial third of the posterosuperior quadrant (mean: 34.4  $\pm$ 5.5 ms versus  $31.1 \pm 3.9$  ms; mean difference: 3.1; 95% CI, 6.2–0.1; p = 0.039) (3a). The ratio was increased in the lateral third (mean:  $1.00 \pm 0.12$  versus  $0.90 \pm 0.15$ ; mean difference: 0.10; 95% CI, 0.18-0.2; p = 0.018) (3b). ICC and RMS-CV were 0.965 and 4% (intraobserver), 0.953 and 4% (interobserver), and 0.988 (all p < 0.001) and 9% (inter-MR scan), respectively.

Conclusions Cartilage T1p MRI mapping in hips is feasible at 1.5 T with strong inter-, intraobserver, and inter-MR scan reliability. The six ROIs (Method 3) showed a difference of T1p values anterolateral quadrant, consistent with the dominant area of cartilage injury in cam femoroacetabular impingement, and antero- and posteromedial, indicating involvement of the entire hip cartilage health. The six ROIs (Method 3) have been shown feasible to assess cartilage damage in hips with a cam deformity using T1p MRI. We suggest applying this ROI selection for further studies using quantitative MRI for assessment of cartilage damage in hips with a cam deformity to achieve better comparability and reproducibility between different studies. The application of this ROI selection on hips with other deformities (eg, pincer deformity, developmental dysplasia of the hip, and acetabular retroversion) has to be analyzed and potentially adapted.

Level of Evidence Level III, diagnostic study.

## Introduction

Cam-type femoroacetabular impingement (FAI) is a common hip deformity and major cause of osteoarthritis (OA) in young adults [10]. Detection of early-stage OA is essential to determine when best to intervene to increase the likelihood that the treatment will decrease pain and improve function [8, 28]. Conventional radiography and standard MRI are insensitive for early biochemical cartilage damage. Advanced MRI techniques have emerged for detection and quantification of biochemical changes in cartilage, even before macroscopic damage occurs. Biochemical changes in early-stage OA include loss of proteoglycan content [6]. One technique, which is sensitive to the proteoglycan content in cartilage, is delayed gadolinium-enhanced MRI of cartilage (dGEMRIC). It has been shown that dGEMRIC can serve in the evaluation of hips with dysplasia [22] or a cam deformity [4]. However, dGEMRIC requires intravenous injection of an exogenous contrast agent; other quantitative approaches for cartilage imaging that do not use contrast agents now are available, including T1p mapping. T1p relaxation time has been shown to correlate inversely with cartilage proteoglycan concentration in bovine and human cartilage samples, making it a potentially useful tool for detection of early biochemical changes in cartilage tissue occurring in the OA disease [27, 31]. Several studies have demonstrated the feasibility of T1p mapping to assess cartilage condition in the knee [11, 13, 21, 23] and hip [26, 29].

The pattern of cartilage damage in hips is influenced by the underlying pathology. Beck et al. [3] first described intraoperative cartilage damage in hips with cam-type FAI mainly in the anterosuperior quadrant, which is consistent with other intraoperative observations [2, 14, 20] and findings in computer simulation [30]. Although MRI mapping of cartilage in the hip has been shown to be feasible and capable to evaluate the biochemical status of cartilage in dysplasia [19, 25, 26, 29] as well as cam FAI (Table 1) [4, 7, 9, 19, 24–26, 29], the region-of-interest (ROI) selection used in terms of size and location has varied between studies. Having said that, understanding how to best define and analyze the ROIs is still evolving and is critical to ensure reproducible and comparable results between institutions and studies.

Therefore, the first aim of this study was to determine representative ROIs for cartilage T1 $\rho$  mapping in hips with a cam deformity. The second aim was to assess intra- and interobserver reliability for cartilage T1 $\rho$  mapping in hips with a cam deformity.

## **Patients and Methods**

This prospective, comparative study was approved by the local ethics committee with informed consent obtained. Fifty-eight patients (58 hips) were enrolled; 38 asymptomatic patients were part of a previous study [12] and 20 patients with unilateral cam-type FAI were recruited in the institution. The study, including the 38 asymptomatic patients, analyzed the prevalence of cam deformity in asymptomatic volunteers. T1 $\rho$  values in these patients have not been analyzed and published previously.

Four MRIs were excluded as a result of motion artifacts. The final study group consisted of 54 hips (44% right) in 54 patients with an average age of  $35 \pm 7$  years; 88% were men (Table 2). MRIs were acquired between December 2010 and November 2013.

Table 1. Regions of interest (ROIs) in	quantitative MRI ar	nalyzing hip	cartilage
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Reference	Region	Zones		Slices			Femoral/ cartilage	acetabular	Total ROI	MRI	
	(o'clock)	Number	Interval (°)	Number	Location	Thickness (mm)	Separate	Combined		T1p	dGEMRIC
Subburaj et al. [29]	4:00-7:00	9	30	1	Central	4	х	х	9	x	
Rakhra et al. [25]	3:00-9:00	4	45	7	Lateral $\rightarrow$ medial	3		х	28	х	
McGuffin et al. [19]	3:00-9:00	2	90	7	Lateral $\rightarrow$ medial	3		х	14	х	
Bittersohl et al. [4]	5:00-9:00	8	30	2	$1 \times lateral/1 \times central$			х	16		x
Domayer et al. [7]	4:00-8:00	8	30	2	$1 \times lateral/1 \times central$			x	16		х
Pollard et al. [24]	3:00-12:00	1	90	1	Lateral	9		x	1		x

The table displays studies analyzing cartilage in hips with cam morphology with advanced MRI techniques (T1 $\rho$  and dGEMRIC) using different ROIs; dGEMRIC = delayed gadolinium-enhanced MRI of cartilage.

Table 2. Demographics of the study group

Parameters	All hips	Hips with a cam deformity	Control hips	p value
Number of hips (number of subjects)	54 (54)	38 (38)	16 (16)	_
Age (years)	35 ± 7 (23–51)	35 ± 8 (23–51)	34 ± 7 (23–47)	0.381
Sex (% male)	88	89	87	0.582
Side (% right)	44	42	50	0.367

The study group was divided into hips with a cam deformity and control hips; there was no difference in age, sex, or side of the included hip between; values of continuous parameters are expressed as mean  $\pm$  SD with range in parentheses.

Two groups were defined: 38 hips with a cam deformity and 16 control hips. A cam deformity was defined as an alpha angle  $> 51^{\circ}$  at the 3:00 position and/or  $> 60^{\circ}$  at the 1:30 position [1]. Of the 38 patients with a cam deformity, 20 were pain-free and 18 symptomatic patients underwent surgery after 6 months of failed nonsurgical management of antiinflammatories and physical therapy.

Exclusion criteria were radiologic signs of osteoarthritis with Tönnis grade of 2 or higher as well as previous hip surgery.

Between the two groups, there was no difference in age or gender (Table 2). None of the patients had an underlying hip disease, previous surgery, or radiologic signs for OA.

T1p imaging was performed on a 1.5-T MRI scanner (Magnetom Symphony; Siemens Medical Solutions, Erlangen, Germany) using the built-in body coil for signal transmission and a flexible four-channel surface coil for signal reception. Five different T1p-weighted data sets (spin-lock times [TSL] = 12, 18, 25, 35, and 45 ms) were acquired in a sagittal oblique plane using a standard turbo spin echo sequence combined with a spin-lock preparation module [20, 25] with the following parameters: spin-lock amplitude (B<sub>1</sub>) = 400 Hz, repetition time = 274 ms, echo time = 13 ms, field of view =  $18 \times 18 \text{ cm}^2$ , matrix =  $384 \times 384$ , in-plane resolution =  $0.46 \times 0.46 \text{ mm}^2$ , number of

slices = 22, slice thickness = 3 mm, number of averages = 1, and total scan time = 21 minutes.

The data sets were postprocessed using custom programs written in MATLAB (MathWorks Inc, Natick, MA, USA). Three-millimeter thick slices, starting from the lateral rim of the acetabular sourcil and progressing medially, were selected from the first (TSL = 12 ms) T1 $\rho$ -weighted data set (Fig. 1A). T1 $\rho$  relaxation maps were calculated by fitting the T1 $\rho$ -weighted images pixelwise to a monoexponential decay function with two unknown parameters (S<sub>0</sub> and T1 $\rho$ ) using the Levenberg-Marquardt algorithm:

 $S(TSL) = S_0 \cdot exp(-TSL/T1\rho),$ 

where S is the signal intensity and  $S_0$  is the signal intensity at TSL = 0 ms [16].

In the second step the cartilage was segmented on the first T1 $\rho$ -weighted data set (Fig. 1B). Six 30° zones in the sagittal plane were defined semiautomatically (Fig. 1C). Subsequently in each hip, a total of six zones in the sagittal pane was analyzed. Depending on the depth of the acetabulum, between six and nine slices of each 3 mm were segmented in the transverse plane. Therefore, between 36 (six zones in the sagittal plane × six slices in the transverse plane) and 54 ROIs (six zones × nine slices) were assessed.



Fig. 1A–C T1 $\rho$  mapping in hip cartilage is demonstrated. T1 $\rho$ -weighted data sets (A) were processed using custom programs written in MATLAB. The acetabular and femoral cartilage bilayer was manually segmented (B) and semiautomatically divided into six zones (C).



Fig. 2 Six final ROIs in  $T1\rho$  mapping in hip cartilage on an axial slice are shown. The hip cartilage was divided into an anterosuperior and a posterosuperior quadrant and into a lateral, an intermediate, and a medial third. The six colors illustrate the six final ROIs on a axial slice of the hip.

Each ROI contained both the femoral and acetabular cartilage of a 30° wide interval in the sagittal plane and of a 3mm thickness in the transverse plane (Fig. 2).

Three different ROI selection methods were analyzed: (Method 1) mean T1 $\rho$  value of the entire hip cartilage; (Method 2) all 36 to 54 ROIs were individually assessed; and (Method 3) six large ROIs were defined.

#### Method 1

All ROIs segmented in each hip were added to one ROI representing the femoral and acetabular hip cartilage expanding anterior from 3:00 o'clock position to posterior 9:00 o'clock position and reaching lateral from the acetabular rim to the most medial part of the hip cartilage at the same level in a coronal plane as the 3:00 o'clock and 9:00 o'clock positions. The mean T1 $\rho$  relaxation time of the entire hip cartilage was combined based on the number of pixels within the ROIs.

#### Method 2

Each of the initially segmented 36 to 54 small ROIs (sections of  $30^{\circ}$  wide intervals in the sagittal plane and 3-mm slices in the transverse plane) were compared within the two groups.

# Method 3a

The 36 to 54 small ROIs were collapsed to generate six bigger ROIs. This ROI selection was based on intraoperative and CT-based observations [3, 30] describing the maximal cartilage damage at 1:00 clock position, expanding from 12:00 to 2:00. This led to a division into an anterosuperior and a posterosuperior quadrant. Furthermore, the cartilage damage was described with an average width of one-third of the acetabular depth. This led to a division into lateral, intermediate, and medial thirds. The subsequent six ROIs were: the anterosuperior and posterosuperior quadrants, divided into lateral, intermediate, and medial thirds. Each ROI spans 90° in the sagittal plane and one-third of the acetabular depth in the transverse plane. To obtain the anterosuperior quadrant, Zones 1 to 3 were combined; to obtain the posterosuperior quadrant, Zones 4 to 6 were combined. The lateral, intermediate, and medial thirds were combined as follows: In hips with six slices, each third included two slices; in hips with seven slices, the two lateral slices represented the lateral third, the three intermediate slices, the intermediate third, and the two medial slices, the medial third. In hips with eight slices, the three lateral slices represented the lateral third, the two intermediate slices the intermediate third, and the three medial slices the medial third. In hips with nine slices, each third contained three slices.

 $T1\rho$  values in each ROI were normalized based on the number of pixels.

# Method 3b

The ratio of  $T1\rho$  values of the anterosuperior over the posterosuperior quadrant was calculated within each third. The relatively spared posterosuperior quadrant was used as an internal control.

To assess interobserver reliability, 10 patients (five in the control group and five in the test group) were analyzed by two observers (HA, GM). For intraobserver reliability, 20 hip MRIs (10 in the control group and 10 in the test group) were analyzed twice by one observer within a time period of 2 weeks (HA). To assess interscan reliability, three patients underwent two scans within a time period of 2 weeks and were analyzed twice by one observer (HA).

#### Statistical Analysis

A Kolmogorov-Smirnov test was run to confirm normal distribution. T1 $\rho$  values were compared using independent samples Student's t-test. Interclass correlation coefficient (ICC) and root mean square coefficient of variation (RMS-CV) were used to analyze intraobserver, interobserver, and interscan reliability. All statistical analyses were performed with the SPSS<sup>®</sup> statistics software package (Version 22; IBM Corp, Armonk, NY, USA).

## Results

The first study aim was to determine appropriate ROIs for cartilage  $T1\rho$  mapping in hips with a cam deformity. This

each of those methods are summarized in sequence here.

The second study aim was to assess intra- and interobserver reliability for cartilage  $T1\rho$  mapping in hips with a cam deformity; results of those analyses follow the mapping results.

was done using four methods: 1, 2, 3a, and 3b; results using

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Method 1: Entire Cartilage (all zones and slices combined)

The mean T1 $\rho$  value in the entire cartilage in all hips combined was 33 ± 4 ms (range, 27–46 ms).

The mean T1p value in the entire cartilage in hips with cam deformity  $(34 \pm 4 \text{ ms})$  was higher than in control hips  $(31 \pm 3 \text{ ms}; \text{mean difference: } 2.5; 95\% \text{ confidence interval [CI]}, 4.7-0.4; p = 0.019; Table 3).$ 

#### Method 2: Analysis of All Small ROIs

The control hips show a pattern with low T1 $\rho$  values in the anterolateral part and high T1 $\rho$  values in the posterolateral part (Fig. 3A). This pattern is lost in hips with a cam deformity (Fig. 3B). An increase in T1 $\rho$  values in hips with a cam deformity was found in lateral slices two and three and in anterosuperior zones one to three. Furthermore, there was an increase in T1 $\rho$  values found in medial slices six to nine, mainly in anterolateral zones one to three but also in posterior zones five and six (Table 4; Fig. 3C).

Table 3. T1p relaxation times in cartilage in hips with and without cam deformity

Region of interest	Cam deformity group	Control-group	p value
Entire cartilage	34 ± 4 (28–46)	32 ± 3 (27–37)	0.019*
Anterosuperior quadrant			
Lateral	33 ± 5 (24–45)	$29 \pm 4$ (22–36)	0.039*
Intermediate	33 ± 5 (23–44)	31 ± 4 (24–36)	0.150
Medial	37 ± 6 (23–50)	33 ± 4 (26–39)	0.014*
Posterosuperior quadrant			
Lateral	33 ± 4 (27–41)	33 ± 4 (24–39)	0.660
Intermediate	33 ± 4 (26–48)	32 ± 7 (24–40)	0.197
Medial	$34 \pm 6$ (20–52)	31 ± 4 (21–37)	0.039*
Ratio anterosuperior to poste	erosuperior quadrant		
Lateral	$1.00 \pm 0.12 \ (0.77 - 1.33)$	$0.90 \pm 0.15 \ (0.65 - 1.24)$	0.018*
Intermediate	$1.01 \pm 0.14 \ (0.64 - 1.31)$	$1.01 \pm 0.17 \ (0.76 - 1.51)$	0.950
Medial	$1.08 \pm 0.17 \ (0.71 - 1.69)$	$1.06 \pm 0.13 \ (0.89 - 1.35)$	0.678

Values are mean  $\pm$  SD with ranges in parentheses; \* significant difference between hips with and without a cam deformity; T1p values in the entire hip cartilage in the anterosuperior and posterosuperior quadrants are in milliseconds; the quadrants are further divided in lateral, intermediate, and medial thirds.

Fig. 3A-C T1p mapping of cartilage in hips with and without cam morphology showing the difference between the two groups using the original 54 ROIs. The figure illustrates T1p values of cartilage in control hips (A), in hips with cam morphology (B), and the difference between the two groups (C) with a grav-coded scale. The nine rings illustrate slices one to nine (lateral to medial). The six wedges illustrate Zones 1 to 6 (anterior to posterior). The bar on the right side decodes the gray scale to the T1p values (in ms) or to the difference (in percentage) between hips with and without cam morphology.



Method 3a: Six Large ROIs

Control hips showed a pattern with low T1p values in the lateral third of the anterosuperior quadrant (29  $\pm$  4 ms) and moderate T1p values in the remaining five ROIs (between 31 and 33 ms) (Fig. 4A). Hips with a cam deformity showed a pattern with high T1p values in the medial third of the anterosuperior quadrant  $(37 \pm 6 \text{ ms})$  and moderate T1p values in the remaining five ROIs (between 32 and 34 ms) (Fig. 4B). Subsequently, comparing hips with and without a cam deformity, T1p values were increased in the lateral and medial thirds in the anterosuperior quadrant (mean:  $32.3 \pm 4.9$  ms versus  $29.4 \pm 4.1$  ms; mean difference: 3.0; 95% CI, 5.8–0.2; p = 0.039 and mean 36.5  $\pm$ 5.6 ms versus 32.6  $\pm$  3.8 ms; mean difference: 3.8; 95% CI, 6.9-0.8; p = 0.014) and in the medial third in the posterosuperior quadrant (mean:  $34.4 \pm 5.5$  ms versus 31.1 $\pm$  3.9 ms; mean difference: 3.1; 95% CI, 6.2–0.1; p = 0.039) (Table 3; Fig. 4C).

Method 3b: Ratio Analysis

The ratio of the anterosuperior to the posterosuperior quadrant was increased in the lateral third (mean:  $1.00 \pm 0.12$  versus  $0.90 \pm 0.15$ ; mean difference: 0.10; 95% CI, 0.18-0.2; p = 0.018), but not in the intermediate and medial thirds in hips with a cam deformity compared with control hips (Table 3).

Intraobserver, Interobserver, and Inter-MR Scan Assessment

The T1p mapping and analysis protocol resulted in strong intraobserver, interobserver, and inter-MR scan reliability. ICC and RMS-CV were 0.965 (p < 0.001) and 4% (intraobserver), 0.953 (p < 0.001) and 4% (interobserver), and 0.988 (p < 0.001) and 9% (inter-MR scan), respectively.

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Slice Zone	1 (lateral)	2	3	4	5	6	7	8	9 (medial)
T1ρ values in	control hips								
1 (anterior)	$25 \pm 5 \ (16 - 33)$	$26 \pm 5 \; (19 - 36)$	$26 \pm 6 \; (15 - 38)$	$28 \pm 5 \ (19 - 38)$	$30 \pm 6 \ (20-41)$	32 ± 6 (23-42)	32 ± 4 (28-42)	$30 \pm 3 \ (26 - 34)$	$28 \pm 5 \ (16 - 38)$
2	$28 \pm 5 \; (16 - 38)$	29 ± 7 (16–42)	$30 \pm 7 \; (18-41)$	31 ± 7 (19-42)	33 ± 4 (26–47)	$34 \pm 6 \ (26-47)$	$32 \pm 5 \ (20 - 39)$	32 ± 4 (25–39)	$33 \pm 2 \ (30 - 34)$
3	$31 \pm 4 \ (24 - 37)$	31 ± 4 (22–37)	$30\pm 5~(20-36)$	$30 \pm 5 \ (21 - 37)$	32 ± 5 (18-38)	32 ± 6 (23-40)	32 ± 7 (21–44)	32 ± 6 (24-43)	$32 \pm 5 \ (26 - 37)$
4	$34 \pm 5 \ (24-47)$	33 ± 4 (25–38)	$32 \pm 5 \ (23 - 38)$	$31 \pm 4 \ (22 - 37)$	$31 \pm 6 \ (21 - 39)$	$31 \pm 5 \ (21 - 38)$	$30 \pm 5 \ (27-43)$	33 ± 5 (27–43)	35 ± 9 (29–49)
5	33 ± 5 (24-42)	33 ± 7 (21–44)	32 ± 7 (19–45)	33 ± 5 (23-43)	$32 \pm 6 \ (23-46)$	32 ± 4 (27–42)	$31 \pm 3 \ (26 - 36)$	$31 \pm 5 \ (22 - 39)$	$32 \pm 5 \ (29 - 36)$
6 (posterior)	35 ± 8 (30–47)	$43 \pm 9 \ (29-52)$	34 ± 8 (20-46)	$30 \pm 6 \; (19-42)$	33 ± 7 (20-48)	31 ± 8 (18-46)	$30 \pm 6 \; (18-43)$	28 ± 9 (12-37)	30 (30)
T1ρ values in	hips with cam defo	armity							
1 (anterior)	30 ± 8 (16-52)	$30 \pm 7 \; (18 - 50)$	$32 \pm 6 \ (19-47)$	32 ± 7 (20-48)	33 ± 8 (20-49)	$35 \pm 7 \ (20-49)$	35 ± 7 (19–48)	37 ± 7 (25–56)	$39 \pm 6 \ (25-43)$
2	32 ± 6 (18-46)	33 ± 7 (23-50)	33 ± 7 (20–47)	33 ± 6 (19-45)	35 ± 6 (23-48)	$36 \pm 6 \ (23-48)$	$37 \pm 7 \ (20 - 50)$	$40 \pm 6 \; (28 - 51)$	$43 \pm 6 \ (35-54)$
3	33 ± 5 (22–47)	33 ± 5 (24-46)	$34 \pm 5 (23 - 43)$	33 ± 5 (25–47)	$34 \pm 5 (25-45)$	$35 \pm 5 \ (21-60)$	$36 \pm 7 \ (21 - 60)$	$38 \pm 5 (32 - 51)$	$42 \pm 7 \ (35-51)$
4	33 ± 5 (23-43)	32 ± 5 (25-45)	32 ± 4 (23-42)	33 ± 5 (21-48)	33 ± 5 (21-52)	33 ± 6 (21-52)	33 ± 7 (20-60)	$34 \pm 7 \ (14-55)$	$43 \pm 10 \; (32-60)$
5	32 ± 5 (22-43)	33 ± 5 (23-44)	33 ± 5 (18-44)	33 ± 5 (23-42)	35 ± 7 (16–56)	$36 \pm 5 \ (25-52)$	$36 \pm 6 \ (23-56)$	35 ± 6 (23-45)	$39 \pm 4 \ (34-43)$
6 (posterior)	$28 \pm 9 \ (9-44)$	$31 \pm 9 \ (15-51)$	33 ± 9 (16-52)	31 ± 7 (16-52)	31 ± 7 (16–48)	33 ± 7 (19-50)	33 ± 7 (16-50)	35 ± 7 (28–50)	37 ± 6 (32-46)
p values									
1 (anterior)	0.071	0.061	0.001*	0.081	0.109	0.189	0.283	0.003*	0.033*
2	0.051	0.038*	0.112	0.389	0.308	0.310	0.017*	0.000*	$0.022^{*}$
3	0.218	0.177	0.029*	0.085	0.083	0.053	0.079	0.002*	0.060
4	0.587	0.706	0.977	0.235	0.289	0.161	0.162	0.479	0.306
5	0.316	0.857	0.997	0.611	0.313	0.023*	$0.013^{*}$	0.075	0.135
6 (posterior)	0.215	0.065	0.565	0.764	0.980	0.230	0.200	$0.014^{*}$	0.384
Comparison o parenthesis; *	f T1 $\rho$ values in hips a p value < 0.05 w	with and without ca as considered signif	um deformity was ca ficant; ROIs = regio	alculated using indep ons of interest.	pendent Student's t-	test; T1ρ values are	in milliseconds and	presented as mean :	± SD with range in

Table 4. T1p relaxation time in cartilage in hips with cam-type deformity versus cartilage in control hips, initial 54 ROIs

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Fig. 4A-C T1p mapping of cartilage in hips with and without cam morphology showing the difference between the two groups using the final six ROIs. The figure illustrates T10 values of cartilage using the final six ROIs in control hips (A), in hips with cam morphology (B), and the difference between the two groups  $(\mathbf{C})$  with a grav-coded scale. The hip cartilage was divided into an anterosuperior and posterosuperior quadrant and in the lateral (lat), intermediate (int), and the medial (med) thirds. The bar on the right side decodes the gray scale to the T1p values (in ms) or to the difference (in percentage) between hips with and without cam morphology.



#### Discussion

Radiography and standard MRI are unable to assess earlystage, microscopic cartilage damage before macroscopic damage occurs. Over the last several years, numerous imaging modalities such as T1 $\rho$ , T2, dGEMRIC, diffusionweighted imaging, sodium imaging, and gagCEST MRI have evolved [18]. These techniques have been applied mostly to the knee, because the hip has always represented a greater challenge as a result of its anatomic features (ie, sphericity).

The aims of this study were to determine representative ROIs for cartilage T1 $\rho$  mapping in hips with a cam deformity and to assess intra- and interobserver reliability for cartilage T1 $\rho$  mapping in hips with a cam deformity. The six ROIs (Method 3) were based on described intraoperative findings and showed a difference of T1 $\rho$  values in this study between hips with and without a cam deformity and are recommended for further studies analyzing T1 $\rho$  MRI in cartilage in hips with a cam deformity. Strong inter-, intraobserver, and inter-MR scan reliability was found for T1 $\rho$  imaging and analysis of the hip cartilage at 1.5 T.

A limitation of the study is that femoral cartilage and acetabular cartilage were assessed as a combined bilayer. With the spatial resolution used in the study, it was not possible to confidently distinguish between the two layers. We suspect that some patients with a cam deformity and advanced cartilage damage have falsely low T1 $\rho$  values as a result of loss of acetabular cartilage; subsequently, the T1 $\rho$  values in these hips predominantly reflect the cartilage status of the remaining cartilage on the femoral head, which is less exposed to damage in hips with cam deformity. Having said that, Subburaj et al. [29], using 3-T MRI, showed that despite using 3 T, the bilayer technique is actually more reproducible than the single-layer acetabular cartilage and equal to a femoral head cartilage single layer.

In this study, our first method analyzed T1 $\rho$  values in the entire hip cartilage and the second T1 $\rho$  values were divided into 36 to 54 ROIs, depending on the size of the hip. Each ROI contained the femoral and acetabular cartilage, a 30° wide interval in the sagittal plane and was 3 mm thick in the transverse plane. In control hips, a distinctive pattern with low T1 $\rho$  values in the anterolateral part has been previously been reported [25]. In comparison, hips with a cam deformity showed increased T1 $\rho$  values mainly in ROIs in the lateral and medial slices in Zones 1 to 3.

Our second method, the assessment of 36 to 54 individual ROIs, was based on the number of available slices. This partition led to increased spatial information, but also to a large number of ROIs, which makes a comparison. especially between different hip sizes, difficult. Subsequently, we strategically collapsed the original small ROIs to six larger and distinct ROIs, reflecting the known anatomic regions of the hip preferentially damaged in cam FAI seen at surgery [3] mainly in the lateral third of the cartilage and in the anterosuperior quadrant. The difference in the 36 to 54 ROIs was mainly seen in the lateral third of the anterosuperior quadrant and in the medial third (more in the anterosuperior than in the posterosuperior quadrant). When collapsing this into six ROIs, prolonged T1p relaxation time was found in the lateral and medial thirds of the anterosuperior quadrant, consistent with the pathomechanism of cam FAI. The same areas showed a difference in T1p values by using the method with 36 to 54 ROIs or the method with six ROIs. Subsequently, it can be said that collapsing the ROIs does not lead to a loss of significance with the added advantage of not being influenced by the size of the hip. The ratio between the anterosuperior and the posterosuperior quadrants was used to eliminate intrinsic interpatient variability. Similar ratios using uninvolved hip cartilage as an intrinsic control have been proposed previously [15, 19, 24]. A difference was found in the ratio in the lateral third, the area exposed to the impingement.

Increased T1 $\rho$  values in the entire hip cartilage and particularly in the anterosuperior area were consistent with the findings published by Subburaj et al. [29]. Mamisch et al. [17], who used dGEMRIC in hips and described decreased T1 values in hips with cam-type deformity in the entire cartilage but especially in the anterosuperior quadrant, indicated loss of proteoglycan mainly in this region. Interestingly, the T1 $\rho$  prolongation was not only found in the lateral third of the anterosuperior cartilage, where the cam impingement occurs and the cartilage damage is expected, but also in the medial third in both the anterosuperior and posterosuperior quadrants. This indicates the degeneration of the cartilage occurs in areas without direct impingement by the cam deformity.

The T1 $\rho$  mapping and analysis protocol developed and used in this study resulted in high intra- and interobserver ICC and low RMS-CV. Similar findings were published in other studies analyzing T1 $\rho$  in the hip cartilage [5, 29] indicating strong intra- and interobserver reliability.

Cartilage T1 $\rho$  MRI mapping in hips is feasible at 1.5 T. The six ROIs (Method 3) have been shown feasible to assess cartilage damage in hips with a cam deformity using

T1p MRI. We suggest applying this ROI selection for further studies using quantitative MRI for assessment of cartilage damage in hips with a cam deformity to achieve better comparability and reproducibility between different studies. The application of this ROI selection on hips with other deformities (eg, pincer deformity, developmental dysplasia of the hip, and acetabular retroversion) has to be analyzed and potentially adapted.

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