## UCSF

UC San Francisco Previously Published Works

## Title

A novel mr-based method for detection of cartilage delamination in femoroacetabular impingement patients

Permalink https://escholarship.org/uc/item/3911k1p7

Journal Journal of Orthopaedic Research®, 36(3)

ISSN

0736-0266

Authors

Samaan, Michael A Pedoia, Valentina Zhang, Alan L <u>et al.</u>

Publication Date 2018-03-01

DOI 10.1002/jor.23667

Peer reviewed



# **HHS Public Access**

Author manuscript *J Orthop Res.* Author manuscript; available in PMC 2019 March 01.

Published in final edited form as:

J Orthop Res. 2018 March ; 36(3): 971–978. doi:10.1002/jor.23667.

# A Novel MR-based Method for Detection of Cartilage Delamination in Femoroacetabular Impingement Patients

Michael A. Samaan<sup>1</sup>, Valentina Pedoia<sup>1</sup>, Alan L. Zhang<sup>2</sup>, Matthew C. Gallo<sup>1</sup>, Thomas M. Link<sup>1</sup>, Richard B. Souza<sup>1,3</sup>, and Sharmila Majumdar<sup>1</sup>

<sup>1</sup>Department of Radiology and Biomedical Imaging, University of California – San Francisco, San Francisco, CA

<sup>2</sup>Department of Orthopaedic Surgery, University of California-San Francisco, San Francisco, CA

<sup>3</sup>Department of Physical Therapy and Rehabilitation Science, University of California-San Francisco, San Francisco, CA

### Abstract

In this study, quantitative magnetic resonance based measurements were used to evaluate  $T_{10}$  and  $T_2$  mapping and heterogeneity in femoroacetabular impingement (FAI) patients with acetabular cartilage delamination and to determine the ability of these quantitative MR-based measurements in detecting delamination. Unilateral hip joint MR-scans of 36 FAI patients with arthroscopicallyconfirmed acetabular cartilage delamination and 36 age, gender and BMI matched controls were obtained.  $T_{1p}$  and  $T_2$  mapping and heterogeneity of the hip joint articular cartilage were assessed in both groups using voxel-based relaxometry (VBR). Quantitative MR-based measurements were compared using statistical parametric mapping (SPM). Receiver operating characteristic (ROC) analysis was used to assess the ability of these quantitative measurements in detecting delamination by calculating the area under the curve (AUC). Pearson partial correlations (r) were used to assess for associations between T<sub>1p</sub> and T<sub>2</sub> radial heterogeneity with the alpha angle in FAI patients. T<sub>10</sub> and T<sub>2</sub> global acetabular values were significantly higher in FAI patients with a focal increase within the posterior acetabular cartilage. FAI patients exhibited increased anterior superior acetabular  $T_{10}$  and  $T_2$  heterogeneity and both of these measures demonstrated a strong ability to detect acetabular cartilage delamination (T1p AUC: 0.96, p<0.001; T2 AUC: 0.93, p<0.001). FAI patients with a larger alpha angle exhibited increased anterior superior acetabular  $T_{1\rho}$  (r=0.48, p=0.02) and  $T_2$  (r=0.42, p=0.03) heterogeneity.  $T_{1\rho}$  and  $T_2$  heterogeneity within the anterior superior acetabular cartilage was shown to be a sensitive measure in detecting

- Conception and design: Michael A Samaan and Valentina Pedoia
- Collection and processing of data: Michael A Samaan, Valentina Pedoia and Matthew C. Gallo
- Analysis and interpretation of the data: All authors
- Drafting of the article: Michael A Samaan and Valentina Pedoia
- Obtaining funding: Alan L. Zhang, Sharmila Majumdar and Richard Souza
- Final approval of the article: All authors

**Corresponding Author:** Michael A. Samaan, Department of Radiology and Biomedical Imaging, University of California-San Francisco, 185 Berry Street, Lobby 6, Suite 350, San Francisco, CA 94107, Telephone: (415) 514-8266, michael.samaan@ucsf.edu. **Authors' Contributions:** 

delamination and may prove beneficial to clinicians in determining optimal interventions for FAI patients.

#### Keywords

Voxel Based Relaxometry (VBR); Radial Heterogeneity; Femoroacetabular Impingement; Delamination;  $T_{1\rho}/T_2$ 

#### INTRODUCTION

Femoroacetabular impingement (FAI) is a morphological disorder of the hip joint, which causes abnormal joint loading patterns and may cause acetabular cartilage delamination<sup>1–4</sup>. Delamination is a separation of the acetabular cartilage layer from the subchondral bone<sup>3,5,6</sup> and is strongly associated with the cam-type deformity<sup>1–4</sup>. More specifically, it was shown that FAI patients with an alpha angle of at least 65° have a 4-fold increase in risk of acetabular cartilage delamination<sup>2</sup>. This injury pattern is ascribed to increased shear forces within the hip joint, as a result of the geometric impingement of the cam lesion. Chronic impingement may then cause enlargement of the cartilage flap and lead to complete detachment from the adjacent cartilage thereby producing loose bodies, full-cartilage thickness defects, and eventually progressive osteoarthritis<sup>1,3,6,7</sup>. Delamination occurs in approximately 30 – 52% of FAI patients<sup>1,8,9</sup> and is usually detected arthroscopically, as delamination is difficult to diagnose radiologically<sup>6,10</sup>. An improved method of detecting acetabular cartilage delamination in FAI patients prior to surgery would benefit the orthopaedic community and may aid in improving surgical treatment options<sup>9,11</sup>.

MR-arthrograms are considered the gold standard in detecting labral abnormalities in FAI patients<sup>12,13</sup> yet radiological detection of acetabular cartilage delamination may still be difficult for radiologists to detect even with MR-arthrograms<sup>1,6,10</sup>. This may be due to the limited amount of distensibility in the hip joint<sup>10</sup>, thereby causing difficulty in viewing the separation between the acetabular and femoral cartilage surfaces. Previous work by Pfirrmann *et al* (2008) demonstrated that when using MR arthrography in detecting acetabular cartilage delamination, fluid under the delaminated cartilage is a specific yet uncommon sign of delamination, while a hypo-intense signal in the acetabular cartilage may be a useful sign to help diagnose cartilage delamination<sup>9</sup>. Overall, radiologists encounter difficulty in assessing delamination due to various factors and therefore, a more quantitative and precise method of detecting delamination through non-invasive imaging methods would benefit the orthopaedic community.

Quantitative MR-imaging (QMRI) techniques such as  $T_{1\rho}$  and  $T_2$  mapping have gained increased use in assessing hip joint cartilage composition in FAI patients<sup>14–17</sup>. Voxel based relaxometry (VBR) allows for the examination of the local distribution of  $T_{1\rho}$  and  $T_2$ mapping and has recently been used to assess both hip<sup>18</sup> and knee<sup>19,20</sup> joint cartilage composition. VBR analysis of the acetabular and femoral cartilage layers of the hip joint was shown to be consistent with a traditional region of interest (ROI) based method in those with hip osteoarthritis (OA)<sup>18</sup> and was shown to be more sensitive in detecting local patterns of  $T_{1p}$  elevation compared to a traditional ROI based assessment<sup>19</sup>. This suggests that VBR

may provide for a more sensitive ability to detect localized differences in articular cartilage composition in FAI patients as well. Also, it is important to understand the effects of FAI on spatial distribution (heterogeneity) of cartilage relaxation times, using texture based analysis, as previous studies have shown that the spatial distribution of cartilage relaxation times may be a sensitive enough measure to discriminate between those with and without  $OA^{21-23}$ .

Therefore, the aims of our study were to: 1) assess the global and local characteristics and heterogeneity of biochemical composition within the acetabular and femoral cartilage using absolute  $T_{1\rho}$  and  $T_2$  VBR and standard deviation of the VBR mapping in subjects with and without FAI, 2) determine whether these quantitative MR-based measurements are sensitive enough to identify acetabular cartilage delamination and 3) determine whether there is an association between acetabular cartilage  $T_{1\rho}$  and  $T_2$  heterogeneity with the cam-type morphology in FAI patients with delamination. We hypothesized that when compared to healthy controls,  $T_{1\rho}$  and  $T_2$  cartilage mapping and heterogeneity within the anterior acetabular cartilage  $T_{1\rho}$  and  $T_2$  heterogeneity within the anterior acetabular cartilage  $T_{1\rho}$  and  $T_2$  heterogeneity within the atterior acetabular cartilage  $T_{1\rho}$  and  $T_2$  heterogeneity would be able to reliably detect delamination and would be associated with the cam-type morphology.

#### METHODS

#### **Patient Demographics**

This case-control study (Level III) was approved by the Committee for Human Research at our institution. All subjects provided written informed consent prior to testing. A total of 72 study participants were enrolled in the current study: 36 FAI patients with acetabular cartilage delamination and 36 age, gender and body mass index (BMI) matched controls were recruited for the current study. Acetabular cartilage delamination was verified arthroscopically during surgical treatment of FAI patients (Figure 1).

All FAI patients were recruited from the Sports Medicine Clinic at our institution and were tested (approximately one month) prior to hip arthroscopy. Each FAI patient exhibited an alpha angle >  $55^{\circ 24}$  and/or a lateral center edge (LCE) angle >  $35^{\circ 25}$  as well as other morphological abnormalities including cartilage and labral abnormalities, osseous bump formation or acetabular over-coverage. The alpha and lateral center edge angles were measured on oblique axial MR-images and standard anterior-posterior pelvis radiographs, respectively. Each FAI patient also demonstrated a positive flexion adduction internal rotation (FADIR) test<sup>26</sup> upon examination by an orthopaedic surgeon (A.L.Z.). Participants were excluded from this study if they possessed a BMI > 35kg·m<sup>-2</sup>, previous hip surgery on the affected side, radiographic indication of osteoarthritis (Kellgren-Lawrence grade<sup>27</sup> > 1 and less than 2mm of joint space) or contraindications to MRI (i.e. pregnancy, coronary stent, etc.). It should be noted that all control participants demonstrated negative FADIR tests, indicating no clinical signs of impingement.

#### **MR-Image Acquisition**

Each study participant underwent a unilateral MR-scan, using a 3 Tesla MR scanner (GE MR750, GE Healthcare, Waukesha, WI) and an 8-channel cardiac coil (GE Healthcare,

Waukesha, WI). For all scans, each participant's feet were aligned in neutral rotation and secured in place to reduce hip movement. The MRI protocol included morphological sequences consisting of  $T_2$  fast spin echo (FSE) images and a combined  $T_{1\rho}$  and  $T_2$  sequence<sup>28,29</sup>, which were used to assess morphological abnormalities and cartilage relaxation mapping, respectively.

The T<sub>2</sub> intermediate-weighted, fat-suppressed, FSE images were obtained with a repetition time of 2400–3700ms, echo time of 60ms, field of view (FOV) of 14 – 20cm, matrix size of 288 × 224 and a slice thickness of 3 – 4mm. These T<sub>2</sub> FSE images were obtained in the sagittal, oblique axial and coronal planes and were used to assess for cartilage and labrum abnormalities, osseous bump formations and acetabular over-coverage in all study participants. In the current study, T<sub>1p</sub> and T<sub>2</sub> MR-images were obtained using the following sequence parameters: FOV of 14cm, matrix size of 256 × 128 pixels, slice thickness of 4mm, recovery time of 1.2s, views per segment of 64, bandwidth of 62.5kHz, no gap, inplane spatial resolution of 0.5mm, in-plane pixel spacing is [0.5469,0.5469] mm and the out of plane slice thickness is 4 mm, time spin lock of 0/15/30/45ms, spin lock frequency of 300Hz and an acquisition time of 13:47.

#### **MR-Image Processing and Analysis**

A previously published method used to implement VBR to assess hip joint cartilage composition<sup>18</sup> was used in the current study and is briefly explained below. Each study participant's MR-data was registered to a single reference participant's MR-data, which provided for the comparison of similar anatomic areas across all participants. The reference data was selected through an iterative process that minimized the global deformation (i.e. minimum deformation template) across participants. Morphological differences across the participants required the use of non-rigid registration applied using the Elastix Toolbox<sup>30</sup>. Non-rigid registration was performed using an iterative process that aimed in reducing the overall differences in femoral head shape between the reference image and in each participant's image. The femoral head center in the reference image and in each participant's image was determined using the Hough transform<sup>31,32</sup>, a method of object detection based on the transformation of all points of an object as defined by a specific equation. For the current study, the femoral head was modeled as a circle, which represented a section of the femoral head in 2D space. The deformation field computed on the first T10-weighted image (TSL=0) was then applied to all the later  $T_{1\rho}$ -weighted and  $T_2$ -weighted images, allowing for the fitting of T<sub>1p</sub> and T<sub>2</sub> maps aligned to the template of each subject. Acetabular and femoral cartilage ROIs manually defined on the reference image were then applied to all the deformed cases creating a fully automatic segmentation procedure.

A radial local standard deviation filter was applied on the  $T_{1\rho}$  and  $T_2$  maps to compute local heterogeneity. We modeled the 2D coronal section of the femoral head as a circle. Prior to the application of the kernel, the 3D  $T_{1\rho}$  and  $T_2$  maps were converted into cylindrical coordinates [ $\rho$ ,  $\theta$ , z] (polar coordinates in the coronal plane) and the kernel applied as a standard deviation filter was [3×1], with 1 in  $\theta$  and 3 in the  $\rho$  direction and followed the circular shape of the femoral head within the 2D section. After the filter's application, each voxel assumes the value of the standard deviation of the [3×1] voxel neighborhood. Radial

symmetry padding was used for the voxels lying on the cartilage borders. This filtering technique returned maps that represent the  $T_{1\rho}$  and  $T_2$  local radial heterogeneity.

#### Statistics

Group demographics, alpha and lateral center edge angles were compared using independent t-tests, where a p-value of less than 0.05 was considered significant. Global differences in  $T_{1\rho}$  and  $T_2$  mapping and heterogeneity between the FAI and control groups in both the acetabular and femoral cartilage layers were assessed using independent t-tests. VBR based group comparisons were performed using Statistical Parametric Mapping (SPM), which provided an assessment of the local differences in  $T_{1\rho}$  and  $T_2$  mapping and heterogeneity between both groups. Group comparisons of the VBR measurements (i.e. mean and standard deviation) were performed using independent t-tests and used to obtain the volumetric p-value SPMs. Also, the percent of significant voxels (PSV) and the average p-value within the significant cluster (p < 0.05) of the acetabular and femoral cartilage layers were computed. Random field theory was used to account for any false positives due to multiple comparisons<sup>33</sup>.

The ability of the global and local  $T_{1\rho}$  and  $T_2$  mapping and heterogeneity in discriminating between FAI patients with delamination and healthy controls was assessed using the area under the curve (AUC) of the receiver operating characteristic (ROC) analysis. The maxefficiency cut-off values were also determined for each of the global and local  $T_{1\rho}$  and  $T_2$  MR-based parameters. The significance of the discriminations (p < 0.05) was assessed using one-tailed independent t-tests.

The association between alpha angle with  $T_{1\rho}$  and  $T_2$  mapping and heterogeneity within the anterior superior acetabular cartilage of the FAI patients was assessed using a Pearson partial correlation coefficient (r), adjusting for age, gender and BMI, where a p-value of less than 0.05 was considered significant. The Jarque-Bera test was performed to check for normality of the  $T_{1\rho}$  and  $T_2$  data distribution<sup>34</sup>.

#### RESULTS

As the FAI patients and healthy controls were matched for age, gender and BMI, these demographics along with the alpha and lateral center edge angles of all study participants were found to be similar (p > 0.05) (Table 1).

The FAI patients exhibited significantly higher global acetabular  $T_{1\rho}$  values (FAI: 34.5±4.0 ms, Control: 32.8±2.6 ms, p = 0.04) yet no differences were exhibited in global femoral  $T_{1\rho}$  values (FAI: 34.7±3.6 ms, Control: 34.8±3.5 ms, p = 0.93) (Figure 2A). More specifically, the FAI patients demonstrated a significant  $T_{1\rho}$  elevation within the posterior acetabulum (Figure 2B; PSV: 8.64%; p = 0.03). The FAI patients exhibited increased global  $T_{1\rho}$  heterogeneity within both the acetabular (FAI: 9.81±2.03 ms, Control: 8.39±1.54 ms, p = 0.001) and femoral (FAI: 8.66±1.46 ms, Control: 7.57±1.46 ms, p = 0.002) cartilage layers (Figure 2C). When assessing the local  $T_{1\rho}$  heterogeneity, the FAI patients exhibited increased  $T_{1\rho}$  heterogeneity within the anterior-superior acetabular (PSV: 10.6%; p = 0.02)

and anterior femoral (PSV: 10.5%; p = 0.02) cartilage layers (Figure 2D). All global and local based QMRI measurements are summarized in Table 2.

Similar to the global  $T_{1p}$  results, the FAI patients exhibited significantly higher global acetabular  $T_2$  values (FAI: 28.8±4.6 ms, Control: 26.5±2.2 ms, p = 0.01) and similar global femoral  $T_2$  values (FAI: 31.3±4.1 ms, Control: 30.9±2.5 ms, p = 0.65) (Figure 3A). The increased acetabular  $T_2$  values occurred in the posterior region of the acetabular cartilage (Figure 3B; PSV: 14.7%; p = 0.02). Global  $T_2$  heterogeneity in the FAI patients was increased in both the acetabular (FAI: 10.1±1.93 ms, Control: 7.91±1.79 ms, p < 0.001) and femoral (FAI: 8.54±2.13 ms, Control: 6.88±1.95 ms, p = 0.001) cartilage layers (Figure 3C). Within both the anterior superior acetabular and anterior femoral cartilage layers, local  $T_2$  heterogeneity was increased and demonstrated a similar amount of significantly different voxels within these regions (PSV=24%, p = 0.02) when compared to healthy controls (Figure 3D).

The ROC analysis (Figure 4) demonstrated that the local heterogeneity of  $T_{1\rho}$  (cut-off: 8.96ms, AUC: 0.96, p<0.001) and  $T_2$  (cut-off: 8.16ms, AUC: 0.93, p<0.001) within the anterior-superior acetabular cartilage better discriminated those with delamination compared to any of the other QMRI based measurements (Table 3).

Within the FAI patients, a larger alpha angle was associated with an increase in  $T_{1\rho}$  (r = 0.48, p = 0.02) and  $T_2$  (r = 0.42, p = 0.03) radial heterogeneity within the anterior superior acetabular cartilage. Although both the  $T_{1\rho}$  and  $T_2$  radial heterogeneity displayed moderate associations with the alpha angle,  $T_2$  radial heterogeneity demonstrated a slightly larger amount of significant voxels that were associated with the alpha angle within FAI patients ( $T_{1\rho}$  PSV: 2.05%,  $T_2$  PSV: 3.38%). The correlation map between  $T_2$  radial heterogeneity and alpha angle in the FAI patients, along with the scatterplot of the average  $T_2$  radial heterogeneity and alpha angles within the anterior superior acetabular cartilage are shown in Figure 5.

#### DISCUSSION

The results of the current study demonstrated that a VBR analysis of quantitative MR-based measurements was able to reliably detect acetabular cartilage delamination within FAI patients. More specifically,  $T_{1\rho}$  and  $T_2$  radial heterogeneity were able to better detect delamination compared to global  $T_{1\rho}$  and  $T_2$  mapping. These results suggest that  $T_{1\rho}$  and  $T_2$  radial heterogeneity may be a more sensitive measure than global  $T_{1\rho}$  and  $T_2$  mapping in detection of delamination. In addition, the  $T_{1\rho}$  and  $T_2$  radial heterogeneity within the anterior superior acetabular cartilage layer, the area of the hip joint in which delamination occurs, was moderately associated with the cam impingement (i.e. alpha angle) and helps to validate previous work suggesting that there is an association between the cam impingement and delamination<sup>1-4</sup>.

In the current study, both the acetabular and femoral global  $T_{1\rho}$  and  $T_2$  relaxation times in the FAI patients were similar to previously published  $T_{1\rho}$  and  $T_2$  relaxation times in FAI patients<sup>14,15</sup>. The FAI patients in the current study exhibited significantly higher global

Samaan et al.

acetabular  $T_{1\rho}$  and  $T_2$  relaxation times compared to the healthy controls with no differences in femoral  $T_{1\rho}$  and  $T_2$  relaxation times. These results indicate altered acetabular cartilage composition and are supported by previous work that demonstrated a strong association between acetabular cartilage abnormalities and increased hip joint loading during gait in FAI patients<sup>35</sup>. These FAI patients may potentially be applying larger loads within the hip joint and these loads seem to be affecting the acetabular cartilage composition more so than the femoral cartilage composition. The increase in T<sub>1p</sub> and T<sub>2</sub> relaxation times may be driven by a more focal difference in the T<sub>10</sub> and T<sub>2</sub> mapping of the posterior acetabular cartilage, where approximately 8 - 14% of voxels within the posterior acetabular cartilage layer demonstrated significantly increased T<sub>10</sub> and T<sub>2</sub> relaxation times in FAI patients when compared to healthy controls. The altered cartilage composition present in the posterior acetabulum of FAI patients may be due to increased shear stresses present within this region of the acetabular cartilage during activities of daily living such as a squatting task<sup>36</sup> and may be indicative of the contre-coup lesion<sup>3,37</sup>. Future studies combining quantitative MRimaging and finite element analyses may provide an ability to understand the mechanical loading patterns during squatting, walking, etc. that may lead to altered hip joint cartilage composition within FAI patients.

Both global acetabular and femoral T<sub>10</sub> and T<sub>2</sub> radial heterogeneity are increased in FAI patients compared to healthy controls but more specifically, the FAI patients in the current study exhibited significantly increased T<sub>1p</sub> and T<sub>2</sub> radial heterogeneity within the anterior superior acetabular cartilage corresponding to the area where delamination is typically observed<sup>9</sup>. Although there were no differences in local  $T_{10}$  and  $T_2$  mapping within the anterior superior acetabular cartilage, the results of the current study suggest that  $T_{10}$  and  $T_2$ radial heterogeneity within the anterior superior region may be more affected by the FAI pathology and/or the presence of cartilage delamination. It is plausible that the shear forces applied to the anterior superior acetabular cartilage by the cam lesion, the same force thought to lead to delamination<sup>3</sup>, may cause disruption of the extracellular matrix architecture in FAI patients. Also, in FAI patients with a cam lesion, provocative motions into hip flexion and internal rotation increases during both static and dynamic activities and the largest pressures and stresses are observed in the anterior superior acetabular cartilage<sup>36,38</sup>. Over time, these increased stresses applied to the anterior superior acetabular cartilage may cause a disruption in the  $T_{1\rho}$  and  $T_2$  spatial distribution of neighboring voxels (increased heterogeneity) and may indicate cartilage damage<sup>21</sup>.

ROC analysis demonstrated that the  $T_{1\rho}$  and  $T_2$  radial heterogeneity within the anterior superior acetabular cartilage layer were able to better detect delamination compared to any of the other  $T_{1\rho}$  and  $T_2$  based parameters analyzed in this study. The anterior superior acetabular  $T_{1\rho}$  and  $T_2$  radial heterogeneity demonstrated similar AUC values ( $T_{1\rho}$ : 0.96,  $T_2$ : 0.93) and suggest that both of these parameters may be useful in detecting acetabular cartilage delamination. Also, the results of the current study help to support the notion that cam impingement is associated with acetabular cartilage delamination<sup>1-4</sup>. More specifically, the FAI patients with worse cam impingement (higher alpha angle) exhibited increased anterior superior acetabular  $T_{1\rho}$  and  $T_2$  radial heterogeneity. Local voxel based assessment of the  $T_{1\rho}$  and  $T_2$  radial heterogeneity exhibited that  $T_2$  radial heterogeneity showed a larger percentage of voxels ( $T_{1\rho}$  PSV: 2.05%,  $T_2$  PSV: 3.38%) within the anterior superior

Samaan et al.

acetabular cartilage that were moderately correlated with the alpha angle compared to  $T_{1\rho}$  radial heterogeneity. These results may suggest that  $T_2$  radial heterogeneity may be slightly more affected by the cam lesion that leads to acetabular cartilage delamination.

The limitations of the current study should be considered when interpreting the findings. The global and local T<sub>1p</sub> and T<sub>2</sub> mapping and heterogeneity were assessed within the central portion of the hip joint and not in the medial and lateral portions, due to the curved surface of the hip joint, which causes challenges in image processing in the far medial and lateral surfaces of the joint<sup>39</sup>. The relatively low resolution of the T<sub>10</sub> and T<sub>2</sub> sequences used in this study (pixel spacing  $0.5459 \times 0.5459$  mm<sup>2</sup>) needs to be acknowledged as a study limitation. The average cartilage plate for the reference hip consists of 4.41 voxels, thereby limiting the number of standard deviation observations made along the radial direction. In this study, we used the assumption of a circular shape for the reference hip, while the registration of the entire dataset on a single template make this assumption true for all the subjects, future work should consider more complex flattening techniques to model the local variation in the curvature of the femoral head surface. These analyses were only performed on FAI patients with arthroscopic validation of delamination and a sample of control subjects who were assumed to have normal hip joint cartilage. A similar study should be performed in FAI patients without delamination that undergo hip femoroplasty, in order to understand the potential differences in hip joint cartilage radial heterogeneity between FAI patients with and without delamination. Also, it is unknown when delamination occurs in these FAI patients and therefore, it is difficult to suggest how radial heterogeneity within the anterior superior cartilage may differ as an effect of time in FAI patients with delamination. In addition, the cross-sectional nature of this study allows for the establishment of associations between MR-based measures and delamination yet a prospective approach is required to help establish the causation of delamination and whether or not MR-based measures, such as  $T_{10}$ and T<sub>2</sub> heterogeneity, would aid in determining early markers of delamination.

In conclusion, quantitative MR-imaging and use of voxel-based relaxometry provide a useful and beneficial assessment of hip joint cartilage composition that may be associated with FAI. More so, using these techniques to quantify the radial heterogeneity within the anterior superior acetabular cartilage was shown to be associated with radiological measurements (alpha angle) and may help clinicians better assess the possibility of an FAI patient exhibiting acetabular cartilage delamination. The approach used in this study can potentially be applied in a clinical setting and may help clinicians and surgeons in determining optimal surgical treatment options for FAI patients.

#### Acknowledgments

This study was supported by the NIH/NIAMS P50 AR060752, F32 AR069458 and the American Orthopaedic Society for Sports Medicine (AOSSM) Young Investigator Grant (YIG-2016-1). The content of this study is solely the responsibility of the authors and does not necessarily represent the views of the NIH or AOSSM.

#### References

 Anderson LA, Peters CL, Park BB, et al. Acetabular cartilage delamination in femoroacetabular impingement. Risk factors and magnetic resonance imaging diagnosis. J Bone Joint Surg Am. 2009; 91:305–313. [PubMed: 19181974]

- Beaule PE, Hynes K, Parker G, et al. Can the alpha angle assessment of cam impingement predict acetabular cartilage delamination? Clin Orthop Relat Res. 2012; 470:3361–3367. [PubMed: 23001504]
- 3. Beck M, Kalhor M, Leunig M, et al. Hip morphology influences the pattern of damage to the acetabular cartilage: Femoroacetabular impingement as a cause of early osteoarthritis of the hip. J Bone Joint Surg Br. 2005; 87:1012–1018. [PubMed: 15972923]
- Johnston TL, Schenker ML, Briggs KK, et al. Relationship between offset angle alpha and hip chondral injury in femoroacetabular impingement. Arthroscopy. 2008; 24:669–675. [PubMed: 18514110]
- Peters CL, Erickson J. The etiology and treatment of hip pain in the young adult. J Bone Joint Surg Am. 2006; 88(Suppl 4):20–26. [PubMed: 17142432]
- Beaule PE, Zaragoza E, Copelan N. Magnetic resonance imaging with gadolinium arthrography to assess acetabular cartilage delamination. A report of four cases. J Bone Joint Surg Am. 2004; 86-a: 2294–2298. [PubMed: 15466743]
- Beaule PE, Zaragoza EJ. Surgical images: Musculoskeletal acetabular cartilage delamination demonstrated by magnetic resonance arthrography: Inverted "oreo" cookie sign. Can J Surg. 2003; 46:463–464. [PubMed: 14680355]
- Fontana A, Mancini D, Gironi A, et al. Hip osteochondral lesions: Arthroscopic evaluation. Hip Int. 2016; 26(Suppl 1):17–22. [PubMed: 27174061]
- 9. Pfirrmann CW, Duc SR, Zanetti M, et al. MR arthrography of acetabular cartilage delamination in femoroacetabular cam impingement. Radiology. 2008; 249:236–241. [PubMed: 18682585]
- Schmid MR, Notzli HP, Zanetti M, et al. Cartilage lesions in the hip: Diagnostic effectiveness of MR arthrography. Radiology. 2003; 226:382–386. [PubMed: 12563129]
- Peters CL, Erickson JA. Treatment of femoro-acetabular impingement with surgical dislocation and debridement in young adults. J Bone Joint Surg Am. 2006; 88:1735–1741. [PubMed: 16882895]
- Leunig M, Werlen S, Ungersbock A, et al. Evaluation of the acetabular labrum by MR arthrography. J Bone Joint Surg Br. 1997; 79:230–234. [PubMed: 9119848]
- Petersilge CA, Haque MA, Petersilge WJ, et al. Acetabular labral tears: Evaluation with MR arthrography. Radiology. 1996; 200:231–235. [PubMed: 8657917]
- Samaan MA, Zhang AL, Gallo MC, et al. Quantitative magnetic resonance arthrography in patients with femoroacetabular impingement. J Magn Reson Imaging. 2016; 44:1539–1545. [PubMed: 27192497]
- Subburaj K, Valentinitsch A, Dillon AB, et al. Regional variations in MR relaxation of hip joint cartilage in subjects with and without femoralacetabular impingement. Magn Reson Imaging. 2013; 31:1129–1136. [PubMed: 23684960]
- 16. Anwander H, Rakhra KS, Melkus G, et al. T1rho hip cartilage mapping in assessing patients with cam morphology: How can we optimize the regions of interest? Clin Orthop Relat Res. 2016
- Rakhra KS, Lattanzio PJ, Cardenas-Blanco A, et al. Can T<sub>1</sub>-rho MRI detect acetabular cartilage degeneration in femoroacetabular impingement?: A pilot study. J Bone Joint Surg Br. 2012; 94:1187–1192. [PubMed: 22933489]
- Pedoia V, Gallo MC, Souza RB, et al. Longitudinal study using voxel-based relaxometry: Association between cartilage T<sub>1p</sub> and T<sub>2</sub> and patient reported outcome changes in hip osteoarthritis. J Magn Reson Imaging. 2017; 45:1523–1533. [PubMed: 27626787]
- Pedoia V, Li X, Su F, et al. Fully automatic analysis of the knee articular cartilage T<sub>1p</sub> relaxation time using voxel-based relaxometry. J Magn Reson Imaging. 2016; 43:970–980. [PubMed: 26443990]
- 20. Russell C, Pedoia V, Amano K, et al. Baseline cartilage quality is associated with voxel-based  $T_{1\rho}$  and  $T_2$  following ACL reconstruction: A multicenter pilot study. J Orthop Res. 2016
- 21. Joseph GB, Baum T, Alizai H, et al. Baseline mean and heterogeneity of MR cartilage T<sub>2</sub> are associated with morphologic degeneration of cartilage, meniscus, and bone marrow over 3 years data from the Osteoarthritis Initiative. Osteoarthritis Cartilage. 2012; 20:727–735. [PubMed: 22503812]

Samaan et al.

- 22. Carballido-Gamio J, Stahl R, Blumenkrantz G, et al. Spatial analysis of magnetic resonance  $T_{1\rho}$  and  $T_2$  relaxation times improves classification between subjects with and without osteoarthritis. Med Phys. 2009; 36:4059–4067. [PubMed: 19810478]
- 23. Li X, Pai A, Blumenkrantz G, et al. Spatial distribution and relationship of T<sub>1p</sub> and T<sub>2</sub> relaxation times in knee cartilage with osteoarthritis. Magn Reson Med. 2009; 61:1310–1318. [PubMed: 19319904]
- Domayer SE, Ziebarth K, Chan J, et al. Femoroacetabular cam-type impingement: Diagnostic sensitivity and specificity of radiographic views compared to radial MRI. Eur J Radiol. 2011; 80:805–810. [PubMed: 21074343]
- Philippon MJ, Ejnisman L, Ellis HB, et al. Outcomes 2 to 5 years following hip arthroscopy for femoroacetabular impingement in the patient aged 11 to 16 years. Arthroscopy. 2012; 28:1255– 1261. [PubMed: 22560486]
- Philippon MJ, Maxwell RB, Johnston TL, et al. Clinical presentation of femoroacetabular impingement. Knee Surg Sports Traumatol Arthrosc. 2007; 15:1041–1047. [PubMed: 17497126]
- Kellgren JH, Lawrence JS. Radiological assessment of osteo-arthrosis. Ann Rheum Dis. 1957; 16:494–502. [PubMed: 13498604]
- Li X, Wyatt C, Rivoire J, et al. Simultaneous acquisition of T1ρ and T2 quantification in knee cartilage: Repeatability and diurnal variation. J Magn Reson Imaging. 2014; 39:1287–1293. [PubMed: 23897756]
- Wyatt C, Kumar D, Subburaj K, et al. Cartilage T1ρ and T2 relaxation times in patients with mildto-moderate radiographic hip osteoarthritis. Arthritis Rheumatol. 2015; 67:1548–1556. [PubMed: 25779656]
- Klein S, Staring M, Murphy K, et al. Elastix: A toolbox for intensity-based medical image registration. IEEE Trans Med Imaging. 2010; 29:196–205. [PubMed: 19923044]
- Tsuji S, Matsumoto F. Detection of ellipses by a modified Hough transformation. IEEE Trans Comput. 1978; C-27:777–781.
- Duda RO, Hart PE. Use of the Hough transformation to detect lines and curves in pictures. Commun ACM. 1972; 15:11–15.
- Marchini J, Presanis A. Comparing methods of analyzing fMRI statistical parametric maps. Neuroimage. 2004; 22:1203–1213. [PubMed: 15219592]
- 34. Jarque CM, Bera AK. A test for normality of observations and regression residuals. International Statistical Review. 1987; 55:163–172.
- 35. Samaan MA, Schwaiger BJ, Gallo MC, et al. Joint loading in the sagittal plane during gait is associated with hip joint abnormalities in patients with femoroacetabular impingement. Am J Sports Med. 2017; 45:810–818. [PubMed: 28006109]
- 36. Ng KC, Rouhi G, Lamontagne M, et al. Finite element analysis examining the effects of cam fai on hip joint mechanical loading using subject-specific geometries during standing and maximum squat. HSS J. 2012; 8:206–212. [PubMed: 24082862]
- 37. Ganz R, Leunig M, Leunig-Ganz K, et al. The etiology of osteoarthritis of the hip: An integrated mechanical concept. Clin Orthop Relat Res. 2008; 466:264–272. [PubMed: 18196405]
- Jorge JP, Simoes FM, Pires EB, et al. Finite element simulations of a hip joint with femoroacetabular impingement. Comput Methods Biomech Biomed Engin. 2014; 17:1275–1284. [PubMed: 23211051]
- 39. Carballido-Gamio J, Link TM, Li X, et al. Feasibility and reproducibility of relaxometry, morphometric, and geometrical measurements of the hip joint with magnetic resonance imaging at 3T. J Magn Reson Imaging. 2008; 28:227–235. [PubMed: 18581346]



#### Figure 1.

Arthroscopic images of a femoroacetabular impingement patient that underwent hip arthroscopy in which the surgeon is probing the region of delamination (left) and showing the separation of the acetabular cartilage from the subchondral bone (right).

Samaan et al.



#### Figure 2.

 $T_{1\rho}$  relaxation mapping (2A – 2B) and radial heterogeneity (2C – 2D) for the femoroacetabular impingement (FAI) and controls (CONT) groups in the acetabular and femoral cartilage layers. The red and white arrows indicate clusters of voxels with significant elevation in  $T_{1\rho}$  relaxation mapping or radial heterogeneity within the anterior and posterior regions, respectively.

Samaan et al.



#### Figure 3.

 $T_2$  relaxation mapping (3A – 3B) and radial heterogeneity (3C – 3D) for the femoroacetabular impingement (FAI) and controls (CONT) groups in the acetabular and femoral cartilage layers. The red and white arrows indicate clusters of voxels with significant elevation in  $T_2$  relaxation mapping or radial heterogeneity within the anterior and posterior regions, respectively.

Samaan et al.



#### Figure 4.

The receiver operating characteristic (ROC) curves for the  $T_{1\rho}$  (left) and  $T_2$  (right) acetabular (ACE) and femoral (FEM) MR-based measurement.  $T_{1\rho}$  and  $T_2$  radial texture within the anterior superior (Ant. Sup.) acetabular cartilage were found to be the most sensitive predictors of delamination in femoroacetabular impingement patients.

Samaan et al.



#### Figure 5.

The Pearson partial correlation map of the  $T_2$  radial heterogeneity (ms) and alpha angle (degrees) within the FAI patients is shown (left). The average  $T_2$  radial heterogeneity within the anterior cartilage (boxed region) demonstrates a high positive correlation with alpha angle measurements (right).

#### Table 1

Demographics and hip joint morphology (mean±standard deviation) for the femoroacetabular impingement (FAI) patients and healthy controls (CONT).

	FAI (N=36)	CONT (N=36)	p-value
Age (years)	35.7±9.4	39.1±10.2	0.14
Gender (Males:Females)	19:17	19:17	1.00
Body Mass Index (kg·m <sup>-2</sup> )	24.1±3.2	23.8±3.1	0.83
Alpha Angle (Degrees)	55.3±11.4	52.0±11.5	0.23
Center Edge Angle (Degrees)	33.4±7.8	30.9±5.9	0.13

#### Table 2

 $T_{1\rho}$  and  $T_2$  mapping and heterogeneity measurements (ms) in the acetabular (ACE) and femoral (FEM) cartilage layers for controls (CONT) and femoroacetabular impingement (FAI) patients, represented as average(standard deviation). An \* represents statistically significant values (p < 0.05).

		FAI (N=36)	CONT (N=36)	p-value
Τ <sub>1ρ</sub>				
	Global ACE	34.5(4.0)	32.8(2.6)	0.04*
	Global FEM	34.7(3.6)	34.8(3.5)	0.93
	Posterior ACE	35.3(4.2)	31.9(2.7)	< 0.001*
	Radial Heterogeneity ACE	9.81(2.02)	8.39(1.54)	0.001*
	Radial Heterogeneity FEM	8.66(1.46)	7.57(1.46)	0.002*
	Radial Heterogeneity Ant. Sup. ACE	11.2(2.3)	7.12(1.24)	< 0.001*
$T_2$				
	Global ACE	28.8(4.6)	26.5(2.2)	0.01*
	Global FEM	31.3(4.1)	30.9(2.5)	0.65
	Posterior ACE	29.6(5.3)	24.9(2.1)	< 0.001*
	Radial Heterogeneity ACE	10.1(1.93)	7.91(1.79)	< 0.001*
	Radial Heterogeneity FEM	8.54(2.13)	6.88(1.95)	0.001*
	Radial Heterogeneity Ant. Sup. ACE	10.6(2.4)	6.70(1.44)	< 0.001*

Anterior Superior: Ant. Sup.

#### Table 3

Receiver operating characteristic (ROC) analysis of the various  $T_{1\rho}$  and  $T_2$  mapping and heterogeneity measurements in the acetabular (ACE) and femoral (FEM), with the corresponding area under the curve (AUC: average [95%CI]) and max efficiency cut-off values (ms). An \* represents statistically significant values (p < 0.05).

		AUC	Cut-off Value	p-value
Τ <sub>1</sub> ρ				
	Global ACE	0.62[0.49 0.75]	33.3	0.04*
	Global FEM	0.49[0.35 0.62]	34.6	0.43
	Posterior ACE	0.75[0.63 0.86]	33.0	< 0.001*
	Radial Heterogeneity ACE	0.71[0.59 0.83]	8.89	0.001*
	Radial Heterogeneity FEM	0.72[0.60 0.84]	7.91	< 0.001*
	Radial Heterogeneity Ant. Sup. ACE	0.96[0.91 1.00]	8.96	< 0.001*
$T_2$				
	Global ACE	0.66[0.53 0.79]	27.3	0.01*
	Global FEM	0.53[0.39 0.67]	31.8	0.32
	Posterior ACE	0.79[0.70 0.89]	26.1	< 0.001*
	Radial Heterogeneity ACE	0.81[0.70 0.91]	8.91	< 0.001*
	Radial Heterogeneity FEM	0.77[0.65 0.88]	7.69	< 0.001*
	Radial Heterogeneity Ant. Sup. ACE	0.93[0.86 0.99]	8.16	< 0.001*

Anterior Superior: Ant. Sup.