Quantitative Knee Cartilage Measurement at MR Imaging of Patients with Anterior Cruciate Ligament Tear

Kazuki Kato
Department of Health Sciences, Hokkaido University, North-12 West-5, Kita-ku, Sapporo City, 060 0812 Japan.

Tamotsu Kamishima, MD, PhD
Faculty of Health Sciences, Hokkaido University, North-12 West-5, Kita-ku, Sapporo City, 060 0812 Japan.

Eiji Kondo, MD
Department of Clinical Support for Medical Practice, Hokkaido University Hospital, North-14 West-5, Kita-ku, Sapporo City, 060 8648 Japan.

Tomohiro Onodera, MD
Department of Clinical Support for Medical Practice, Hokkaido University Hospital,
North-14 West-5, Kita-ku, Sapporo City, 060 8648 Japan.

Shota Ichikawa, RT

Graduate School of Health Sciences, Hokkaido University, North-12 West-5, Kita-ku, Sapporo City, 060 0812 Japan.

Address correspondence to:

Tamotsu Kamishima, MD, PhD,

Faculty of Health Sciences, Hokkaido University

North-12 West-5, Kita-ku, Sapporo City, 060 0812 Japan.

Phone/FAX; 81-11-706-2824

E-mail: ktamotamo2@yahoo.co.jp
Quantitative Knee Cartilage Measurement at MR Imaging of Patients with Anterior Cruciate Ligament Tear
Abstract

In previous studies, numerous approaches were proposed that assess knee cartilage volume quantitatively by using 3D magnetic resonance (MR) imaging. However, clinical use of these approaches is limited because it is prone to metal artifact in postoperative cases. Our purpose in this study was to validate a method for knee cartilage volume quantification by using conventional MR imaging in patients who underwent anterior cruciate ligament (ACL) reconstruction surgery. The study included 16 patients who underwent MR imaging before and 1 year after ACL reconstruction surgery. Knee cartilage volumes were measured by our computer-based method with use of T1-weighted sagittal images. We classified the cartilage into 8 regions and made comparisons between preoperative and postoperative cartilage volumes in each region. There was a significant difference between preoperative and postoperative cartilage volumes with regard to medial posterior weight-bearing, medial posterior, lateral posterior weight-bearing, and lateral posterior portions (p=0.006, p=0.023, p=0.017 and p=0.002, respectively). These results were consistent with the previous studies showing that knee cartilage loss occurs frequently in these portions due to an anterior subluxation
of the tibia accompanied by ACL tear. With our method, knee cartilage volumes could be quantitatively measured with conventional MR imaging in patients who underwent ACL reconstruction surgery.
Key Words

knee cartilage: anterior cruciate ligament: magnetic resonance imaging
Hyaline cartilage covering the surface of the diarthrodial joint is important for normal joint function and can transmit and distribute pressure without sustaining substantial wear [1,2]. In addition, because articular cartilage provides a nearly frictionless gliding surface, these forces can be transmitted during dynamic joint activity [3–8]. In order to satisfy these complex functional requirements, articular cartilage shows unique morphologic features and biomechanical properties that are unmatched by artificial material [1,2,9,10]. Observing cartilage destruction and loss is important for evaluating osteoarthritis (OA) [11,12].

Magnetic resonance (MR) imaging has been in use for cartilage evaluation for many years because it provides high soft-tissue contrast and spatial resolution. MR imaging enables direct qualitative and quantitative visualization and evaluation of the cartilage [13]. In recent years, some investigators have developed computer-based methods to create 3D cartilage reconstruction from magnetic resonance images [12,14–17]. These methods can be used for assessment of cartilage degeneration and cartilage loss quantitatively by evaluation of cartilage volume, while the methods have high susceptibility to metal artifacts due to their sequence design depending on the gradient echo technique, which is not recommended for quantitative analysis of cartilage in
post-operative status.

Anterior cruciate ligament (ACL) tear leads to anterior subluxation of the tibia with engagement of the anterior femoral condyle against the posterior tibial plateau [18–23]. As a result, cartilage loss occurs in the anterior femoral condyle and posterior tibial plateau [19,21]. In addition, 74% of patients who underwent ACL reconstruction surgery displayed some signs of radiographic OA within 7 years post-surgery. Thus, OA is common in ACL-reconstructed knees [24]. Therefore, our purpose in this study was to verify a quantification method for knee cartilage volumes by using conventional MR imaging in patients who had undergone ACL reconstruction surgery.

Materials and methods

Patients

The study included 16 patients who underwent MR imaging before and 1 year after ACL reconstruction surgery (Tables 1,2). This study was conducted in accordance with the Declaration of Helsinki and its updates and was approved by the local ethics committee. Informed consent was obtained from all of the patients.

MR imaging
All patients in this study group underwent MR imaging, with T1-weighted images obtained before (baseline) and 1 year after (follow-up) ACL reconstruction surgery. MR images were performed with the Achieva 1.5T A-series (Philips) and the Achieva 3.0T TX (Philips). Imaging parameters are summarized in Table 3. In follow-up studies, special care was taken so that the slice position was identical to that of the baseline study.

Image analysis

In this study, we used an original software in which cartilage is manually segmented from MR images taken at baseline and at follow-up, and the interval difference in cartilage volume between baseline and follow-up is displayed with Microsoft Excel and a color map.

The original software was developed with Microsoft Visual C# 2013. The MR images taken at baseline and at follow-up were imported into the software. Then, 3 steps for cartilage segmentation and volumetry were followed.

Step-1: semi-automated bone segmentation for bone area elimination at bone/cartilage interface
The interface between the subchondral bone and its cartilage is a sharp curvilinear line in most cases. Therefore, to eliminate bone pixels from volumetry of the cartilage, all the slices in each MR study were binarized automatically by Otsu's method [25], and fused images of the binary images and the original images were created (Fig.1). Here, the threshold was selected to separate bony structures from the soft tissues including cartilage, a pixel value larger than the threshold was displayed in green, and a pixel value less than the threshold was displayed with that of the original images. The green pixels containing bone area were then automatically extracted from the display. When the bone was not completely displayed in green, we manually eliminated the signal from the bone by mouse dragging operation. Extracting the bone using automatic binarization facilitated easier segmentation of the cartilage boundary for the subsequent processing, thus reducing the time for segmentation without affecting the accuracy of cartilage quantification.

Step-2: manual segmentation of the knee cartilage at its interface to joint space

We manually segmented the articular side of the knee cartilage which we deemed cartilage by mouse dragging operation. The segmented part was displayed in red (Fig.2). During cartilage segmentation, the pixels formally displayed in green were never
repainted as red, so that the bony areas were not added to the knee cartilage volumes. In baseline and follow-up images, we carefully performed this operation only on a slice judged to be the same section by observing between baseline and follow-up slices where articular cartilage is visible. The number of slices used for image analysis therefore resulted in the same for baseline as for follow-up. The total of the measured areas was calculated as volume.

Step-3: automated sub-region (8-regions) segmentation and volume calculation

We divided the segmented cartilage into quarters from the pixel where the cartilage exists at the most anterior position to the pixel at the most posterior position in the slices with the cartilage, and divided these into medial and lateral portions. We defined the 8 regions as lateral anterior, lateral anterior weight-bearing, lateral posterior weight-bearing, lateral posterior, medial anterior, medial anterior weight-bearing, medial posterior weight-bearing, and medial posterior portions (Fig.3.). The way we segmented the cartilage came from the findings of previous studies; ACL tear leads to anterior subluxation of the tibia with impaction of the anterior femoral condyle against the posterior tibial plateau along with cartilage loss in the anterior femoral condyle and posterior tibial plateau [19,21]. The amount of cartilage volume change was obtained by
subtraction of the volume of the baseline from that of the follow-up, and color mapping was performed based on the value of these changes (Fig.4.).

One radiological technologist with three months of experience and enough prior knowledge about the position of the cartilage performed computer-based analysis twice with a half of year interval. In order to examine the validity of the original software, pre- and post-operative knee cartilage was quantified using our software (which includes semi-automatic and manual segmentation) and an image analysis software called ImageJ (National Institutes of Health, Bethesda, MD) in one case (20-year-old, male). In ImageJ, we manually segmented cartilage utilizing the “Polygon selection” tool.

Statistical analysis

SPSS version 22.0 (IBM Corp, New York, NY) for Windows was used for statistical analysis. When we compared the paired samples, normality was tested by the Shapiro-Wilk test and did not find normality; we therefore performed the Wilcoxon signed-rank test, which is a nonparametric test.

Intra-observer agreement was estimated using intra-class correlation coefficients (ICC) employing a one-way random effects model for intra-observer
agreement. ICC values were interpreted as poor agreement for values between 0 and 0.20, fair agreement for values between 0.21 and 0.40, moderate agreement for values between 0.41 and 0.60, substantial agreement for values between 0.61 and 0.80, and almost perfect for values between 0.81 and 1.00 [25]. We compared the time taken per slice between the original software and ImageJ in a Wilcoxon signed-rank test.
Results

There was a significant difference between baseline and follow-up cartilage volumes in the medial posterior weight-bearing, medial posterior, lateral posterior weight-bearing, and lateral posterior portions (p=0.006, p=0.023, p=0.017, and p=0.002, respectively). However, there was no significant difference between baseline and follow-up cartilage volumes in the medial anterior, medial anterior weight-bearing, lateral anterior, and lateral anterior weight-bearing portions (p=0.352, p=0.098, p=0.642, and p=0.602, respectively), as shown in Table 4. A representative case is shown in Fig.5.

Intra-observer agreement of knee cartilage volume in 8 regions was almost perfect (ICC = 0.955; 95% confidence interval [95% CI], 0.943-0.965). Intra-observer agreement for delta values (difference between the baseline and follow-up) was substantial (ICC = 0.803; 95% CI, 0.732-0.857). The knee cartilage volume in the original software/ImageJ was 2108/2038 mm3 (0.094% difference) and 2108/2110 mm3 (3.4% difference) at the baseline and follow-up, respectively. The mean time (± standardized deviation) taken with the original software and ImageJ was 50.9 (± 10.1) and 131 (± 0.000788) seconds, respectively (p = 0.00004).

Discussion
In this study, we evaluated cartilage volumes in 16 patients before and 1 year after ACL reconstruction surgery. There was a significant difference between baseline and follow-up cartilage volumes in each of 4 regions (medial posterior weight-bearing, medial posterior, lateral weight-bearing posterior, and lateral posterior portions). In previous studies, quantitative knee cartilage evaluation was conducted using 3D-MR images [12,14–17], but to the best of our knowledge, there were no studies in which there was a quantitative evaluation of postoperative cartilage volume as in this study. T1-weighted images are used routinely for the analysis of anatomic structures. Moreover, they have an advantage over gradient echo sequences and fat suppression images, being free from postoperative metal artifacts. Therefore, this method could be applied immediately to clinical practice.

Previous investigations reported that ACL tear leads to anterior subluxation of the tibia with impaction of the anterior femoral condyle against the posterior tibial plateau along with cartilage loss in the anterior femoral condyle and posterior tibial plateau [19,21]. Also, previous studies showed that ACL-reconstructed knees had greater contact along the medial ridge of the medial plateau and the posterior aspect of the lateral plateau, compared to healthy contralateral knees and to the knees of healthy persons during exercise [15]. Furthermore, in another study, 74% of the patients who
had undergone ACL reconstruction surgery showed several signs of radiographic OA within 7 years after surgery, indicating that early-onset OA is common in ACL-reconstructed knees [24]. For these reasons, cartilage loss may occur in the anterior femoral condyle and posterior tibial condyle at the time of ACL tear. In this study, the results that there was a significant difference between baseline and follow-up cartilage volumes in the medial posterior weight-bearing, medial posterior, lateral weight-bearing posterior, and lateral posterior portions were consistent with this hypothesis. That may suggest that our newly proposed method could successfully quantify knee cartilage volume.

In this study, we showed that there is significant reduction in the reasonable anatomical location of the knee cartilage in patients who underwent ACL reconstruction surgery. Moreover, the knee cartilage volumes can be evaluated by using the conventional MR images with our software. We consider this is due to careful scan planning in terms of stable slice selection and imaging parameter, including slice thickness and slice gap. Together with compatible results from the data using the free software of ImageJ, we considered our cartilage volumetry was to be valid.

In our study results, the knee cartilage volumes did not show any significant difference in the anterior and anterior weight-bearing portions of both medial and lateral
sides. This may be attributed to the fact that we could not divide segmented cartilage into the femoral side and the tibial side. In our study population, as ACL tears were caused by trauma, cartilage wear was considered to be localized. For more sensitive detection of the localized cartilage loss, measurement of cartilage volume should be performed on the femoral side and the tibial side separately. However, in this study, it was difficult to separate the cartilage into the femoral side and the tibial side on the T1-weighted sagittal images.

We believe that as the original software used in this study is accurate, reproducible, and less time-consuming, it can be applied to the comparison among the procedures of ACL reconstruction. In a previous study, the comparison among the procedures of ACL reconstruction was performed, but the investigations evaluated ACL by arthroscopy, whereas a cartilage evaluation was not done [26]. We considered that adding cartilage evaluation to one of the prognostic evaluations might reveal signs of OA, which is considered to occur frequently in ACL-reconstructed knees.

Our study had several limitations. First, the devices used for MR images differ between inter- and intra- patients due to the retrospective design of this study. However, the MR parameters used in this study did not differ greatly between the devices, and the results of this research implied that cartilage volume may be quantified by use of
routine images, independent of the device. The second limitation was the small number
of patients included in this study. Further study on more patients is needed for
confirmation of the results. Thirdly, although the cartilage region is clearly depicted
using 3D imaging methods such as the 3D-FISP sequence, we have no 3D imaging
methods available for comparison. However, we believe that it is meaningful to
quantitatively evaluate cartilage using 2D-T1w MR images although accuracy for
cartilage volumetry may be inferior to 3D sequences due to the slice gap. Finally,
because our original software was not free from manual segmentation for the most part,
cartilage evaluation could be subjective and time-consuming. Moreover, when
conducting similar research in other institutes, there is a possibility that some variation
may occur depending on the analyst’s skill. Therefore, in order fundamentally to solve
these problems, we need to build a new algorithm for automating our original software.

Conclusion

Knee cartilage volumes could be measured quantitatively with conventional MR
imaging in patients who underwent ACL reconstruction surgery.

Compliance with ethical standards
Conflict of interest

All authors have no conflicts of interest to disclose.

Ethical approval

All procedures in studies involving human participants were performed in accordance with the ethical standards of the Faculty of Health Sciences, Hokkaido University, and with the 1964 Helsinki Declaration and its later amendments or comparable ethical standards.

Informed consent

Informed consent was waived by the ethical committee of Faculty of Health Sciences, Hokkaido University as this was a retrospective study.

References

1. Mow VC, Holmes MH, Lai WM. Fluid transport and mechanical properties of


16. Koo S, Gold GE, Andriacchi TP. Considerations in measuring cartilage thickness


fracture patterns of the knee associated with anterior cruciate ligament tears: assessment

23. Graf BK, Cook DA, De Smet AA, Keene JS. "Bone bruises" on magnetic resonance

Changes After Anterior Cruciate Ligament Reconstruction Using Bone-Patellar
Tendon-Bone or Hamstring Tendon Autografts: A Retrospective, 7-Year Radiographic

25. Landis JR, Koch GG. The measurement of observer agreement for categorical data.
Biometrics. 1977;33:159-74.

Tissue Preservation on Clinical and Arthroscopic Results After Anatomic
Table 1. Characteristics of patients

<table>
<thead>
<tr>
<th></th>
<th>Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Total no. of subjects included</td>
<td>16</td>
</tr>
<tr>
<td>Age. mean (range) in years</td>
<td>30.1 (13-58)</td>
</tr>
<tr>
<td>Sex. female/male</td>
<td>7/9</td>
</tr>
<tr>
<td>Knee. right/left</td>
<td>5/11</td>
</tr>
</tbody>
</table>
Table 2. Athletic activity or situation associated with the injury, and period from injury to surgery

<table>
<thead>
<tr>
<th>Case number</th>
<th>Situation</th>
<th>Days</th>
<th>Sex</th>
<th>Age [years]</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>tumble</td>
<td>756</td>
<td>M</td>
<td>36</td>
</tr>
<tr>
<td>2</td>
<td>rugby</td>
<td>92</td>
<td>M</td>
<td>19</td>
</tr>
<tr>
<td>3</td>
<td>ski</td>
<td>85</td>
<td>M</td>
<td>55</td>
</tr>
<tr>
<td>4</td>
<td>soccer</td>
<td>3626</td>
<td>M</td>
<td>32</td>
</tr>
<tr>
<td>5</td>
<td>basketball</td>
<td>136</td>
<td>F</td>
<td>13</td>
</tr>
<tr>
<td>6</td>
<td>badminton</td>
<td>0</td>
<td>M</td>
<td>19</td>
</tr>
<tr>
<td>7</td>
<td>tumble</td>
<td>139</td>
<td>F</td>
<td>45</td>
</tr>
<tr>
<td>8</td>
<td>gym vault</td>
<td>111</td>
<td>M</td>
<td>18</td>
</tr>
<tr>
<td>9</td>
<td>gym vault</td>
<td>15706</td>
<td>F</td>
<td>58</td>
</tr>
<tr>
<td>10</td>
<td>volleyball</td>
<td>57</td>
<td>F</td>
<td>15</td>
</tr>
<tr>
<td>11</td>
<td>volleyball</td>
<td>1461</td>
<td>F</td>
<td>51</td>
</tr>
<tr>
<td>12</td>
<td>volleyball</td>
<td>100</td>
<td>M</td>
<td>49</td>
</tr>
<tr>
<td>13</td>
<td>volleyball</td>
<td>85</td>
<td>F</td>
<td>16</td>
</tr>
<tr>
<td>14</td>
<td>triathlon</td>
<td>110</td>
<td>M</td>
<td>20</td>
</tr>
<tr>
<td>15</td>
<td>rugby</td>
<td>93</td>
<td>M</td>
<td>19</td>
</tr>
<tr>
<td>16</td>
<td>soccer</td>
<td>24</td>
<td>F</td>
<td>17</td>
</tr>
</tbody>
</table>
Table 3. Acquisition parameters of MR imaging

<table>
<thead>
<tr>
<th>Magnetic Field Strength [T]</th>
<th>1.5</th>
<th>3</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sequence</td>
<td>SE</td>
<td>T1WI</td>
</tr>
<tr>
<td>Section</td>
<td>sagittal</td>
<td></td>
</tr>
<tr>
<td>Slice Thickness [mm]</td>
<td>3</td>
<td></td>
</tr>
<tr>
<td>FA [degree]</td>
<td>90</td>
<td></td>
</tr>
<tr>
<td>NEX</td>
<td>2</td>
<td></td>
</tr>
<tr>
<td>Slice Gap [mm]</td>
<td>0.3</td>
<td></td>
</tr>
<tr>
<td>TR [msec]</td>
<td>500 or 579</td>
<td>700 or 800</td>
</tr>
<tr>
<td>TE [msec]</td>
<td>15</td>
<td>10 or 12</td>
</tr>
<tr>
<td>ETL</td>
<td>3 or 4</td>
<td>3</td>
</tr>
<tr>
<td>BW [Hz]</td>
<td>230 - 259</td>
<td>242 - 255</td>
</tr>
<tr>
<td>Pixel Size [mm]</td>
<td>0.222 × 0.222 - 0.390 ×</td>
<td>0.273 × 0.273 - 0.3125 ×</td>
</tr>
<tr>
<td>No. of Slices</td>
<td>23 or 24</td>
<td>23 or 25</td>
</tr>
</tbody>
</table>
Table 4. Comparison between baseline and follow-up cartilage volume in each region

<table>
<thead>
<tr>
<th>Regions</th>
<th>Baseline</th>
<th>Follow-up</th>
<th>P value (&lt;0.05)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>n</td>
<td>Mean [mm³]</td>
<td>SD [mm³]</td>
</tr>
<tr>
<td>Medial</td>
<td>16</td>
<td>31.8</td>
<td>29.0</td>
</tr>
<tr>
<td>Anterior</td>
<td></td>
<td>203</td>
<td>54.6</td>
</tr>
<tr>
<td>Anterior weight-bearing</td>
<td></td>
<td>309</td>
<td>69.2</td>
</tr>
<tr>
<td>Anterior weight-bearing</td>
<td></td>
<td>321</td>
<td>105</td>
</tr>
<tr>
<td>Posterior</td>
<td></td>
<td>115</td>
<td>96.4</td>
</tr>
<tr>
<td>Anterior weight-bearing</td>
<td></td>
<td>164</td>
<td>74.5</td>
</tr>
<tr>
<td>Anterior weight-bearing</td>
<td></td>
<td>376</td>
<td>96.8</td>
</tr>
<tr>
<td>Lateral</td>
<td></td>
<td>351</td>
<td>107</td>
</tr>
</tbody>
</table>

SD = standard deviation
Fig. 1. Original image (A) and fused image (B)
Fig. 2. Segmentation of cartilage at fused image
Fig. 3. Cartilage classification into 8 regions.

(A) Sagittal MR image of the lateral side of the knee (B) Sagittal MR image of the medial side of the knee. (a) Lateral anterior portion (b) Lateral anterior weight-bearing portion (c) Lateral posterior weight-bearing portion (d) Lateral posterior portion (e) Medial anterior portion (f) Medial anterior weight-bearing portion (g) Medial posterior weight-bearing portion (h) Medial posterior portion
The closer to red, the larger the difference between the follow-up and baseline is to the positive value. The closer to blue, the larger the difference between the follow-up and baseline is to the negative value. Green indicates almost unchanged.

(a) Lateral anterior portion (b) Lateral anterior weight-bearing portion (c) Lateral posterior weight-bearing portion (d) Lateral posterior portion (e) Medial anterior portion (f) Medial anterior weight-bearing portion (g) Medial posterior weight-bearing portion (h) Medial posterior portion
Fig.5. Example of a case with significant cartilage volume loss at multiple cartilage regions.

Sagittal T1-w MR images of knee in 36-year-old man before and 1 year after ACL reconstruction surgery. (A) baseline image. (B) is colored cartilage of (A). (c) is followup image. (D) is colored cartilage of (C). The cartilage volume at baseline (followup) was 70.9 (37.6), 131.3 (43.8), 484.9 (314.0) and 458.0 (313.0) mm$^3$ in anterior, anterior weight-bearing, posterior weight-bearing and posterior at baseline, respectively. These interval changes are difficult to recognize using conventional visual assessment.