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Cartilage Degeneration Post-Meniscectomy Performed for Degenerative Disease versus Trauma: Data from the Osteoarthritis Initiative

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Abstract

Purpose.—To compare the extent of cartilage deterioration in knees with prior meniscal resection related to trauma versus knees with resection related to degenerative disease, and to compare cartilage deterioration in knees with meniscal surgery to knees without meniscal surgery, controlling for prior knee trauma.

Methods and Materials.—In this cross-sectional study, we assessed cartilage deterioration in right knees of Osteoarthritis Initiative participants: (i) with meniscal surgery due to injury (n=79); (ii) matched control knees with a prior injury but without meniscal surgery (n=79); (iii) with meniscal surgery but without preceding injury (n=36); and (iv) matched control knees without meniscal surgery or prior knee injury (n=36). Cartilage composition was measured using T2 measurements derived using semi-automatic cartilage segmentation of the right. Linear regression analysis was used to compare compartmental values of T2 between groups.

Results: Comparing the mean T2 values in surgical cases with and without injury our results did not show significant differences (group i vs iii, p>0.05). However, knees with previous meniscal surgery showed significantly (p<0.001) higher mean T2 values across all compartments (i.e. global T2) when compared to those without meniscal surgery for both knees with a history of trauma (group i vs ii) and knees without prior trauma (group iii vs iv). Similar results were obtained when analyzing the compartments separately.

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Competing interest statement

None of the authors have any financial or other interests related to the manuscript submitted to Skeletal Radiology that might constitute a potential conflict of interest.

Conclusion: Cartilage deterioration, assessed by T2, is similar in knees undergoing meniscal surgery after trauma and for degenerative conditions. Both groups demonstrated greater cartilage deterioration than nonsurgical knees, controlling for prior knee injury.

Keywords

Knee osteoarthritis; meniscal surgery; magnetic resonance imaging; risk factors; biomarkers

INTRODUCTION

The biomechanics of the menisci are a critical component in the functionality of the knee joint and play a crucial role in axial load transmission of the knee [1]. Alterations of the menisci impact biomechanical loading and are a frequent source of pain and disability [2]. In this context, meniscal injuries are one of the most common pathologies of the knee and often require medical attention [3]. In general, meniscal injuries occur over a broad age range. While trauma related meniscal injuries are found more often in young adults, degenerative injuries are known to be more often associated with increasing age and are seen with work-related repetitive activities [4]. Both types of meniscal injury contribute to the risk of knee osteoarthritis (OA) [5–7] and previous studies have questioned the benefit of treating meniscal lesions with arthroscopic surgery compared to nonsurgical treatments [8]. However, arthroscopic partial meniscectomy is a frequently performed treatment of torn menisci, [9] and in the United States, it is one of the most frequent orthopaedic surgical procedure [10].

Based on postoperative outcome scores (Knee Injury and Osteoarthritis Outcome Score – KOOS) previous studies have shown a higher evidence of knee OA following excision of degenerated versus traumatic meniscal tears [11, 12]. Additionally, Zikria et al. found that subsequent joint space narrowing (JSN) progression was greater in knees with meniscal surgery without prior trauma compared to meniscal surgery with prior trauma [13]. This is potentially due to the fact that surgical resection of degenerative lesions may only remove the most current evidence of the disorder [14], while the preexisting degraded cartilage architecture of the osteoarthritic knee persists and continues to worsen [15]. However, no previous studies are available to provide insights into the cartilage composition of patients with previous meniscal surgery compared to knees without surgery and the possible differences in the cartilage composition of patients who underwent meniscal surgery for trauma compared to meniscal surgery for degenerative disorders.

To quantitatively assess articular cartilage composition of the knee, MR-based T2 relaxation time measurements have been demonstrated to be a reliable method to reflect changes of hydration and organization of collagen fibrils in the extracellular matrix of the hyaline cartilage [16, 17]. Thus this imaging technique will also provide pertinent insights into the pathophysiology of the cartilage composition in patients who have undergone meniscal surgical procedures.

The purpose of this study was (i) to analyze differences in the articular cartilage composition, using cartilage T2 relaxation time measurements, between knees with previous meniscal surgery related to trauma and those knees with meniscal surgery related to

degenerative disorders and (ii) to compare cartilage deterioration between knees with meniscal resection and non-surgical control knees frequency matched for sex, age, Kellgren-Lawrence (K/L) grade and BMI.

MATERIAL & METHODS

The Osteoarthritis Initiative Database

Participants in our study were selected from the Osteoarthritis Initiative (OAI) database, a multicenter cohort study of knee osteoarthritis, sponsored by the U.S. National Institutes of Health (NIH). In total, 4796 participants were enrolled from February 2004 to May 2006 and completed as of January 1st 2015, creating a an ethnically diverse cohort of women and men ages 45-79 years (mean age, 61 years) that included participants with symptomatic knee OA and subjects with risk factors for OA but without presenting knee OA symptoms.

Following a nationwide advertising campaign, recruitment and enrollment consisted of an initial eligibility assessment by telephone, a screening clinic visit and finally an enrollment clinic visit at one of the four clinical centers (University of Maryland, Baltimore, Maryland; Memorial Hospital of Rhode Island / Brown University, Pawtucket, Rhode Island; Ohio State University, Columbus, Ohio; University of Pittsburgh, Pittsburgh, Pennsylvania). All clinical enrollment center were supervised by a steering committee, the primary governing body of the study and scientific leadership, to ensure uniform enrollment goals for gender and age strata in the primary subcohorts. Among data of the MRI scans, radiographs and biological specimens, the enrollment visit included a detailed clinical assessment of the subjects' knees, questions about use of medication, questionnaires assessing physical disability (due to knee pain and arthritis), knee pain und function as well as the assessment of risk factors for knee OA (including history of knee injury and knee surgery, abnormal biomechanical stress related to physical activity and obesity). Additionally, the participants were been given a self-administered questionnaire to complete at home that was reviewed at the enrollment clinic. The questionnaire included information on education, medical history, smoking/alcohol and income.

The purpose of the OAI database was to develop a public accessible domain research resource to investigate the role of MRI-based imaging biomarkers in an attempt to better understand the disease onset and ultimately prevent its progression (https://nda.nih.gov/oai/) [18]. All scans were acquired over a nine-year period.

Selection of participants from the OAI

In this retrospective cross-sectional analysis study, participants were eligible for inclusion if they had no to mild radiographic signs of knee OA (Kellgren-Lawrence (K/L) classification of osteoarthritis; grade 0–2) and complete data on BMI, sex, and age at the time of the enrollment visit. Participants were excluded if, during the follow-up visits, they self-reported the diagnosis of rheumatoid arthritis or the use of rheumatoid arthritis medication. Finally, to identify participants who underwent meniscal surgery, we used the OAI self-administered questionnaires from each participant (Medical history; Release Version 0.2.2): Participants were asked if they underwent surgery of the meniscus where they repaired or cut away a torn

meniscus. Additionally, participants who had meniscal surgery related to an injury episode were identified through i.) having a history of knee injury and ii.) participant stating that the meniscectomy was to treat an injury. As provided by the OAI questionnaire workbook, injury was defined as "knee ever injured badly enough to limit ability to walk for at least two days". The second surgical group had no history of knee injury and this group was therefore considered to have degenerative meniscal tears. To ensure accuracy of the acquired data, as provided by the questionnaires, we only selected participants that clearly answered to all relevant questions with yes or no and participants that were uncertain about their medical history (replied with "Not Expected", "Don't know/Unknown/Uncertain", or "Refused") were not included. Additionally, all knees with meniscal surgery identified by self-report were reviewed by a trained musculoskeletal radiologist to confirm the reported postsurgical status (DS). Through identification of the age at the time the injury occurred (selfadministered questionnaire: "how old at injury"), age at the time of the arthroscopy (selfadministered questionnaire: "how old at arthroscopy"), and the age at the time of the MRI scan (age at the time point of enrollment) the corresponding delta () between each episode was determined.

At the time point of enrollment (baseline visit), a total of 115 participants with history of meniscal surgery confirmed by MRI review (case cohort) and 3362 participants without meniscal surgery (control cohort) met the inclusion criteria.

Finally, to examine the association of cartilage degeneration with surgically treated meniscal lesions compared to knees without previous surgery for meniscal lesion we also selected for each surgical group a non-surgical group that had the same self-report history with respect to knee injuries (non-surgical control group without history of knee injury, non-surgical control group with history of knee injury), and frequency matched to each surgical group for age, sex. BMI and KL grade. The matching did not include the meniscal lesion status of the nonsurgical knees and, therefore, was unknown and not taken into account in the selection. The flow chart in Figure 1 illustrates the detailed selection of all participants in this study. All MRI scans and the clinical assessment were performed as part of the OA Initiative.

MR imaging protocol

All scans were acquired using 3.0T MRI systems (Siemens Magnetom Trio; Siemens Healthcare, Erlangen, Germany) with quadrature transmit-receive coils (USA Instruments, Aurora, OH, USA) at one of the four OAI clinical sites. The scan protocol consisted of a 3D dual echo steady-state (DESS) gradient-echo with water excitation (WE) sequence obtained in the sagittal plane (16.3/4.7/25°, TR/TE/flip angle, spatial resolution=0.365mm×0.456mm, slice thickness=0.7mm), a sagittal 2D intermediate-weighted fast spin-echo sequence (TR/TE=3200/30ms, spatial resolution=0.357mm×0.511mm, slice thickness=3.0mm), a coronal 2D proton density fast spin-echo sequence (TR/TE=3700/29ms, spatial resolution=0.365mm×0.456mm, slice thickness=3.0mm), as well as a sagittal 2D multi-slice multi-echo (MSME) spin-echo sequence with a total of seven echo times (TEs 10ms, 20ms, 30ms, 40ms, 50ms, 60ms, 70ms), a repetition time (TR) of 2700 ms, a field of view (FOV) of 120mm, a slice thickness of 3mm (with 0.5 mm gap) and an in-plane spatial resolution of 0.313×0.446 mm².

Quantitative cartilage T2 analysis

All cartilage T2 measurements were acquired using the 2D MSME SE images. Cartilage segmentations were obtained for each compartment separately with the compartments being defined as: patella (PAT), lateral femur (LF), medial femur (MF), lateral tibia (LT), and medial tibia (MT). Due to substantial flow artifacts from the popliteal artery, the trochlear region was excluded from the analysis. The cartilage of each compartment was segmented by a trained researcher (KK). The software used for the T2 analysis was an in-house, spline-based algorithm written in MATLAB (the Mathworks, Natick, Massachusetts) that allows semi-automatic segmentation of each compartment by analyzing the T2 values in a mono-exponential decay model as a fitting function for the signal intensity using 6 echoes (TE 20–70 ms) after excluding the first echo in order to minimize systemic errors and improve signal-to-noise ratio [19]. Cartilage T2 analysis was conducted for the mean T2 across all compartments for the global knee joint as well as for each compartment separately.

Statistical Analysis

Statistical analysis was performed using STATA software (Version 14, College Station, TX: StataCorp LP). A p-value of <0.05 was considered to be statistically significant. Demographic data between the study cohorts were assessed using either Pearson's χ 2-test, one-way ANOVA or independent samples t-test as appropriate. Linear regression models were used to assess the differences in cartilage T2 parameters between the groups for every single compartment, the mean across all compartments as well as separately for the weight bearing compartments (mean of both femoral and tibial compartments). All analyses were adjusted for the common risk factors of knee OA for age, sex, and BMI.

To assess intrareader reproducibilities, coefficients of variation (CV) were calculated on a percentage basis as the root mean square average [20, 21]. Averaged over all compartments, the inter-reader reproducibility for T2 measurements was 0.88%. The CVs for each compartment were 0.85% for PAT, 0.93% for MF, 1.24% for LF, 0.79% for MT, and 0.57% for LT. Overall, the reproducibilities were similar to those reported previously [22, 23].

RESULTS

Participant demographics

The subject characteristics are summarized in Table 1. The average age of all participants (n=230) was 57.9 years (SD \pm 8.9) with a BMI of 27.9 (SD \pm 4.0). One-hundred-fifty-four (67.0%) participants were male. The overall K/L grade distribution showed a higher percentage of participants with K/L 2 (n=128; 55.7%). K/L grades of 0 and 1 showed a similar distribution with n=55 (23.9%) and n=47 (20.4%), respectively. Overall, the four groups (groups (i) and (ii), and groups (iii) and (iv)) were well matched with no significant differences (p>0.05) in age, sex, BMI, and K/L grade.

Differences in clinical parameters between all groups.

Assessing the physical activity, the comparison of all groups did not show statistically significant differences (PASE score, p>0.05). The total Western Ontario and McMaster Universities Osteoarthritis Index (WOMAC) did also not show any show statistically

significant differences between all groups. Looking at the WOMAC pain subscale, however, we observed an overall statistical difference (p=0.022). Participants that underwent degenerative meniscal surgery showed a significantly higher WOMAC pain score when compared to non-surgical controls without preceding knee trauma (p=0.004). However, WOMAC pain scores were not statistically different in participants with traumatic meniscal surgery compared with non-surgical controls with a preceding knee trauma (p=0.415). With respect to the age at the time of the arthroscopy, as expected, traumatic meniscal surgery was performed at a younger age than meniscal surgery performed for degenerative changes and thus MRIs performed in the degenerative cohort were performed more recently. All clinical parameters are shown in Table 1.

Differences in cartilage T2 values between participants with meniscal surgery related to trauma and degenerative disease

Our first analysis focused on participants with meniscal surgery only, to possibly identify differences in the cartilage composition with different surgical indication for meniscal surgery. Overall, comparing participants who underwent meniscal surgery due to an injury with participants who underwent meniscal surgery related to a degenerative tear, the cartilage mean T2 values did not show statistically significant differences between the two groups. For the latter group, higher mean T2 values were seen in the global knee (p=0.731) and in the global weight bearing knee (p=0.731) and in the global weight bearing knee compartments (p=0.354). Both the tibial and the lateral femur compartments also showed higher mean T2 values in participants with degenerative meniscal tears but these results also did not translate into statistical differences (Table 2).

Differences in cartilage T2 values between participants with meniscal surgery and nonsurgical controls

Participants who underwent meniscal surgery in relation to a previous injury showed overall higher mean T2 values when compared to controls without meniscal surgery but with a history of knee injury (Figure 2). Mean T2 values were significantly higher for the global knee (mean T2 effect size differences in ms [95% CI]: 0.94 [0.45,1.44], p<0.001) as well as for the global weight bearing knee compartments (0.97 [0.45,1.48], p<0.001). Higher mean T2 values were also observed for every single compartment, with significantly higher values for the lateral tibia, the medial femur, and the medial tibia (p<0.02) (Table 3).

Participants who underwent meniscal surgery to repair a degenerative meniscal tear showed also overall higher mean T2 values when compared to the control group without meniscal surgery and without history of knee injury (Table 3). Mean T2 values were significantly higher for the global knee as well the global weight bearing knee compartments with 1.35 ms [0.60,2.09] (p<0.001) and 1.56 ms [0.79,2.33] (p<0.001), respectively. Similar to the injury group (as shown in Table 2) mean T2 values were again higher in all separate compartments for participants who underwent meniscal surgery when compared to controls without injury. However, only the lateral femur (1.67 [0.59,2.75], p=0.003) and medial tibia (1.27 [0.19,2.34], p=0.02) showed a statistically significant differences.

DISCUSSION

Comparing cartilage T2 values in participants that underwent meniscal surgery our study did not find differences with respect to whether surgery was to repair an injury or degenerative related meniscal tears. However, when compared to matched non-surgical controls, we found that participants who underwent meniscal surgery showed overall higher mean T2 values. These findings indicate that if patients undergo meniscal surgery, the surgical indication (traumatic vs degenerative) does not significantly affect the degree of cartilage deterioration and seems to be equally the same for both groups. However, compared to non-surgical controls, the risk of developing OA, as suggested by the higher mean T2 values, remains higher for patients that did undergo meniscal surgery.

Knee injuries are common and meniscus lesions are one of the most frequent sources of knee pain and disability [2]. Treatment options for meniscal lesions involve non-operative treatment, meniscal repair, or meniscal resection. The different treatment options need to be critically explored since studies have shown that meniscal lesions, both those treated surgically [14], and those not treated surgically [14], are strong risk factors for developing knee OA. With regard to surgically treated meniscal lesions, this increased risk is due to an altered contact pressure on the cartilage surfaces and other biomechanical and biochemical changes that results from a combination of the damaged meniscus and any associated trauma, and changes induced by surgical intervention.[24]. The long-term outcomes of meniscectomy in particular have been well-studied. In a retrospective study design, Englund et al. evaluated the radiographic and clinical outcome of patients who underwent meniscal resection 15-22 years earlier [5] and found that meniscal resection is associated with a threefold greater risk of later development of knee OA compared to knees without clinical evidence of meniscal lesions. They also showed that the increased risk of knee OA in the surgically treated knees was associated with pre-existing early stage OA and with risk factors common to knee OA, including female sex and obesity. Rongen et al. studied the risk of knee replacement in 335 participants who underwent arthroscopic meniscectomy and found a threefold increased risk for future knee replacement surgery of osteoarthritis knees in patients with meniscectomy [25]. In a prospective, longitudinal study design McNicholas et al. were able to show meniscectomy leads to symptomatic osteoarthritis of the knee later in life 30 [26] and 40 [27] years after surgery was done.

Whereas, previous studies showed that meniscal surgery was associated with an increased risk of developing knee OA, none of these studies examined whether the cartilage composition itself was altered in knees with preceding meniscal surgery or whether cartilage composition was different when surgery was to repair a meniscal injury episode versus a degenerative meniscal tear. To directly assess differences in the cartilage architecture of participants that underwent meniscal surgery we based our analysis on cartilage T2 parameters derived from quantitative T2 relaxation time measurements [16]. Overall, our findings support the results of previous studies on meniscal surgery as a risk factor for knee OA. When compared to a non-surgical control group, participants who underwent meniscal surgery showed significantly higher mean T2 values, indicating an altered cartilage matrix composition with an increased risk for knee OA [28, 29]. It is important to acknowledge that since the nonsurgical control knees in our study included an unknown proportion with

meniscal lesions that were not treated surgically, this finding cannot be attributed to the effect of meniscal surgery, per se. Meniscal lesions that are not treated surgically are extremely common [30] and have a substantially increased risk of developing radiographic and symptomatic OA [14]. Isolating the impact of surgical treatments on outcomes in knees with meniscal lesions requires randomized trials, and to date most studies suggest that clinical and structural outcomes of meniscal lesions are similar regardless of type of treatment [3, 8, 31].

In order to identify a possible differences in cartilage composition related to the surgical indication, we also investigated the cartilageT2 profile of participants who underwent surgery for a trauma related meniscal injury compared to surgery for a degenerative lesion. Previous studies using patient reported outcomes, total knee replacement surgery and radiographic evaluation to determine the outcome of OA have suggested an increased risk of knee OA following resection of degenerated menisci compared to trauma induced tears [11–13]

We addressed this question using cartilage T2 values, a biomarker data that directly reflects possible differences in cartilage composition between the two groups. In contrast to previous studies using other outcome measures, we did not find differences in cartilage T2 values between these two surgical groups. Given the pathognomonic of meniscal tears, due to the preexisting osteoarthritic changes in knee with degenerative meniscal surgery [32], one would expect to find higher T2 values in participants that underwent degenerative meniscal tear surgery and, therefore a more advanced cartilage deterioration, when compared to participants that underwent traumatic meniscal tear surgery.

That we did not find such differences may be explained by the fact that the meniscal surgeries and knee injuries in our study may have occurred several years ago, allowing for the long-term impact of the trauma plus the knee surgery to decrease the differences in cartilage deterioration compared to knees with degenerative lesions. In addition, restricting our sample to knees with KL grades of less than 3 the OA resulted in a similar radiographic OA severity between the two groups.

We acknowledge that our study has several limitations. Firstly, data on meniscal surgery and knee injuries in the OAI were based on self-report and no surgical reports were obtained. The OAI database does not provide detailed information about the type of morphological meniscal damage or type of surgery or on how the knee injury occurred (e.g. sport activity, daily activity, work related) which would potentially be helpful to further categorize participants with a history of knee injury related meniscal surgery. However, to ensure an accurate distinction between trauma and degenerative surgery, we used the OAI self-administered questionnaire that, in addition to the question of knee injury and meniscal surgery, also asked if the surgery was to repair an injury. In addition, a musculoskeletal radiologist reviewed the MRIs of all knees self-reported as having meniscal surgery to confirm their post-surgery status. Nevertheless, misclassification of surgical vs nonsurgical knees and surgery for traumatic vs. degenerative lesions is possible and could influence our results. Secondly, the nonsurgical controls were not selected with respect to whether meniscal lesions were present. The presence of meniscal lesions in a portion of control knees

likely causes us to underestimate the T2 differences between surgically treated knees and normal knees without lesions and to overestimate the differences between surgically treated and non-surgically treated knees with meniscal lesions. Thirdly, with respect to our study participants that did not undergo a MRI scan at the enrollment clinic, a different percentage rate between these two groups was observed, posing a potential bias due to a different study participant attrition. The main causes for participants not undergoing a MRI scan were most likely MRI safety, knee replacements, and participants might have declined the MRI scan due to personal reasons not disclosed by the OAI database. Fourthly, arthroscopy for traumatic meniscal surgery was performed at a younger age than meniscal surgery performed for degenerative changes. Hence, the degenerative cohort received their MRI scans more recently. Finally, we performed a retrospective, cross-sectional analysis in which there may be large variations in the time since injury and surgery. Studies of the outcomes of new meniscal injuries and recent surgeries are needed to better describe the consequences of each.

In conclusion, using quantitative cartilage T2 measurements, our study showed that meniscal tears treated surgically have worse cartilage composition compared to knees without a history of surgery for meniscal tears, supporting studies suggesting that meniscal damage treated with surgery is a risk factor for development of knee OA. However, with respect to indication of surgery, we were able to show that differences in the cartilage matrix were non significant in participants who received surgery related to trauma or degenerative meniscal disease, suggesting that they have a similar impact on post-surgical cartilage health.

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REFERENCES

- 1. Fox AJS, Bedi A, Rodeo SA. The Basic Science of Human Knee Menisci. Sports Health 2012;4:340–351. 10.1177/1941738111429419 [PubMed: 23016106]
- 2. Torres L, Dunlop DD, Peterfy C, et al. The relationship between specific tissue lesions and pain severity in persons with knee osteoarthritis. Osteoarthritis Cartilage 2006;14:1033–1040. 10.1016/j.joca.2006.03.015 [PubMed: 16713310]
- Herrlin S, Hållander M, Wange P, Weidenhielm L, Werner S. Arthroscopic or conservative treatment of degenerative medial meniscal tears: a prospective randomised trial. Knee Surg Sports Traumatol Arthrosc Off J ESSKA 2007;15:393

 –401. 10.1007/s00167-006-0243-2
- Snoeker BAM, Bakker EWP, Kegel CAT, Lucas C. Risk factors for meniscal tears: a systematic review including meta-analysis. J Orthop Sports Phys Ther 2013;43:352–367. 10.2519/ jospt.2013.4295 [PubMed: 23628788]

 Englund M, Lohmander LS. Risk factors for symptomatic knee osteoarthritis fifteen to twenty-two years after meniscectomy. Arthritis Rheum 2004;50:2811–2819. 10.1002/art.20489 [PubMed: 15457449]

- Ding C, Martel-Pelletier J, Pelletier J-P, et al. Meniscal tear as an osteoarthritis risk factor in a largely non-osteoarthritic cohort: a cross-sectional study. J Rheumatol 2007;34:776–784 [PubMed: 17361984]
- Jarraya M, Roemer FW, Englund M, et al. Meniscus morphology: Does tear type matter? A narrative review with focus on relevance for osteoarthritis research. Semin Arthritis Rheum 2017;46:552– 561. 10.1016/j.semarthrit.2016.11.005 [PubMed: 28057326]
- Herrlin SV, Wange PO, Lapidus G, Hållander M, Werner S, Weidenhielm L. Is arthroscopic surgery beneficial in treating non-traumatic, degenerative medial meniscal tears? A five year follow-up. Knee Surg Sports Traumatol Arthrosc Off J ESSKA 2013;21:358–364. 10.1007/s00167-012-1960-3
- Rangger C, Klestil T, Gloetzer W, Kemmler G, Benedetto KP. Osteoarthritis after arthroscopic partial meniscectomy. Am J Sports Med 1995;23:240–244. 10.1177/036354659502300219 [PubMed: 7778712]
- Cullen KA, Hall MJ, Golosinskiy A. Ambulatory surgery in the United States, 2006. Natl Health Stat Rep 2009;1–25
- Englund M, Roos EM, Roos HP, Lohmander LS. Patient-relevant outcomes fourteen years after meniscectomy: influence of type of meniscal tear and size of resection. Rheumatol Oxf Engl 2001;40:631–639
- Azam M, Shenoy R. The Role of Arthroscopic Partial Meniscectomy in the Management of Degenerative Meniscus Tears: A Review of the Recent Literature. Open Orthop J 2016;10:797– 804. 10.2174/1874325001610010797 [PubMed: 28217206]
- Zikria B, Hafezi-Nejad N, Roemer FW, Guermazi A, Demehri S. Meniscal Surgery: Risk of Radiographic Joint Space Narrowing Progression and Subsequent Knee Replacement-Data from the Osteoarthritis Initiative. Radiology 2017;282:807–816. 10.1148/radiol.2016160092 [PubMed: 27697006]
- Englund M The role of the meniscus in osteoarthritis genesis. Med Clin North Am 2009;93:37–43,
 x. 10.1016/j.mcna.2008.08.005 [PubMed: 19059020]
- 15. Noble J, Hamblen DL. The pathology of the degenerate meniscus lesion. J Bone Joint Surg Br 1975;57:180–186 [PubMed: 1173585]
- Link TM, Neumann J, Li X. Prestructural cartilage assessment using MRI. J Magn Reson Imaging 2016. 10.1002/jmri.25554
- Prasad AP, Nardo L, Schooler J, Joseph GB, Link TM. T(1)rho and T(2) relaxation times predict progression of knee osteoarthritis. Osteoarthritis Cartilage 2013;21:69–76. 10.1016/ j.joca.2012.09.011 [PubMed: 23059757]
- Felson DT, Nevitt MC. Epidemiologic studies for osteoarthritis: new versus conventional study design approaches. Rheum Clin North Am 2004;30:783–97, vii. 10.1016/j.rdc.2004.07.005
- Raya JG, Dietrich O, Horng A, Weber J, Reiser MF, Glaser C. T2 measurement in articular cartilage: impact of the fitting method on accuracy and precision at low SNR. Magn Reson Med 2010;63:181–93. 10.1002/mrm.22178 [PubMed: 19859960]
- 20. Glüer CC, Blake G, Lu Y, Blunt BA, Jergas M, Genant HK. Accurate assessment of precision errors: how to measure the reproducibility of bone densitometry techniques. Osteoporos Int J Establ Result Coop Eur Found Osteoporos Natl Osteoporos Found USA 1995;5:262–270
- 21. Stehling C, Baum T, Mueller-Hoecker C, et al. A novel fast knee cartilage segmentation technique for T2 measurements at MR imaging--data from the Osteoarthritis Initiative. Osteoarthritis Cartilage 2011;19:984–989. 10.1016/j.joca.2011.04.002 [PubMed: 21515391]
- 22. Gersing AS, Solka M, Joseph GB, et al. Progression of cartilage degeneration and clinical symptoms in obese and overweight individuals is dependent on the amount of weight loss: 48-month data from the Osteoarthritis Initiative. Osteoarthritis Cartilage 2016;24:1126–1134. 10.1016/j.joca.2016.01.984 [PubMed: 26828356]
- 23. Neumann J, Hofmann FC, Heilmeier U, et al. Type 2 diabetes patients have accelerated cartilage matrix degeneration compared to diabetes free controls: data from the Osteoarthritis Initiative. Osteoarthritis Cartilage 2018;26:751–761. 10.1016/j.joca.2018.03.010 [PubMed: 29605381]

24. Haemer JM, Song Y, Carter DR, Giori NJ. Changes in articular cartilage mechanics with meniscectomy: A novel image-based modeling approach and comparison to patterns of OA. J Biomech 2011;44:2307–2312. 10.1016/j.jbiomech.2011.04.014 [PubMed: 21741046]

- 25. Rongen JJ, Rovers MM, van Tienen TG, Buma P, Hannink G. Increased risk for knee replacement surgery after arthroscopic surgery for degenerative meniscal tears: a multi-center longitudinal observational study using data from the osteoarthritis initiative. Osteoarthritis Cartilage 2017;25:23–29. 10.1016/j.joca.2016.09.013 [PubMed: 27712957]
- 26. McNicholas MJ, Rowley DI, McGurty D, et al. Total meniscectomy in adolescence. A thirty-year follow-up. J Bone Joint Surg Br 2000;82:217–221 [PubMed: 10755429]
- Pengas IP, Assiotis A, Nash W, Hatcher J, Banks J, McNicholas MJ. Total meniscectomy in adolescents: a 40-year follow-up. J Bone Joint Surg Br 2012;94:1649–1654.
 10.1302/0301-620X.94B12.30562 [PubMed: 23188906]
- 28. Liebl H, Joseph G, Nevitt MC, et al. Early T2 changes predict onset of radiographic knee osteoarthritis: data from the osteoarthritis initiative. Ann Rheum Dis 2015;74:1353–9. 10.1136/annrheumdis-2013-204157 [PubMed: 24615539]
- 29. Baum T, Stehling C, Joseph GB, et al. Changes in knee cartilage T2 values over 24 months in subjects with and without risk factors for knee osteoarthritis and their association with focal knee lesions at baseline data from the Osteoarthritis Initiative. J Magn Reson Imaging 2012;35:370–378. 10.1002/jmri.22834 [PubMed: 21987496]
- Englund M, Guermazi A, Gale D, et al. Incidental meniscal findings on knee MRI in middle-aged and elderly persons. N Engl J Med 2008;359:1108–1115. 10.1056/NEJMoa0800777 [PubMed: 18784100]
- 31. Graaf VA van de, Noorduyn JCA, Willigenburg NW, et al. Effect of Early Surgery vs Physical Therapy on Knee Function Among Patients With Nonobstructive Meniscal Tears: The ESCAPE Randomized Clinical Trial. JAMA 2018;320:1328–1337. 10.1001/jama.2018.13308 [PubMed: 30285177]
- 32. Howell R, Kumar NS, Patel N, Tom J. Degenerative meniscus: Pathogenesis, diagnosis, and treatment options. World J Orthop 2014;5:597–602. 10.5312/wjo.v5.i5.597 [PubMed: 25405088]

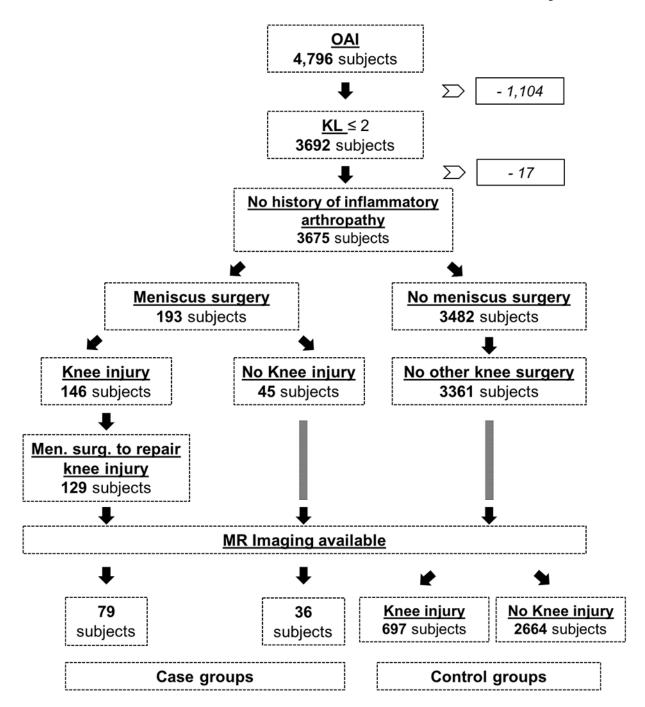


Fig 1. Selection of study participants

Flow-chart illustrating patient selection from the OAI cohort.

No Preceding Trauma

Preceding Trauma

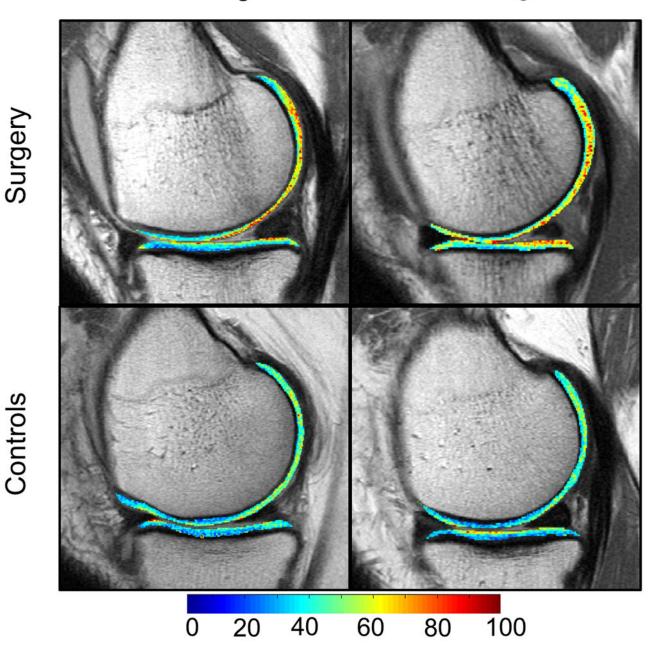


Fig 2. Cartilage T2 color map of participants with and without meniscal surgery
Representative cartilage T2 color maps, overlaid on sagittal 2D multi-slice multi-echo
(MSME) spin-echo sequences, showing the medial knee compartment in each of the
different groups. (a) and (b) show the knee of participants that underwent meniscal surgery
to repair a degenerative (a) and a traumatic tear (b), whereas, (c) and (d) both show
participants without meniscal surgery but with (c) and without (d) preceding trauma. Both
surgical participants show overall higher T2 values (indicated by the yellow and red color
maps) with emphasis on the femoral condyle, including the weight bearing as well as the

posterior aspect of the femoral condyle. The controls show overall predominantly lower T2 values (blue and green color coded).

Table 1.

Participant characteristics

	All ptcp. (n=230)	Traumatic meniscal surgery (n=79)	Non surgery controls with preceding trauma (n=79)	p- value*	Degenerative meniscal surgery (n=36)	Non surgery controls without preceding trauma (n=36)	p- value**	p- value ^{***}
Age	57.9 ± 8.9	56.5 ± 8.7	56.8 ± 8.5	0.860	61.2 ± 9.3	60.3 ± 9.2	0.675	0.013
BMI	27.9 ± 4.0	28.0 ± 4.0	28.4 ± 3.9	0.454	27.5 ± 4.2	26.9 ± 3.9	0.534	0.287
Sex	154 (67.0%) / 76 (33.0%)	57 (72.2%) / 22 (27.8%)	53 (67.1%) / 26 (32.9%)	0.489	24 (66.7%) / 12 (33.3%)	20 (55.6%) / 16 (44.4%)	0.334	0.379
K/L grade				0.979			0.834	0.993
Grade 0	55 (23.9%)	18 (22.8%)	18 (22.8%)		9 (25.0%)	10 (27.8%)		
Grade 1	47 (20.4%)	16 (20.3%)	17 (21.5%)		8 (22.2%)	6 (16.7%)		
Grade 2	128 (55.7%)	45 (57.0%)	44 (55.7%)		19 (52.8%)	20 (55.6%)		
Surgical/injury	characteristics							
Age at MENR		44.9 ± 13.5			54.6 ± 10.7			< 0.001
Age at INJ		43.3 ± 14.3	34.6 ± 15.5	< 0.001				
INJ - MENR		2 ± 6.3						
MENR - MRI		11.6 ± 11.6			6.6 ± 5.4			0.015
Clinical charact	eristics							
PASE	173.8 ± 88.5	180.0 ± 88.6	183.0 ± 94.2	0.835	161.8 ± 82.6	152.5 ± 79.1	0.629	0.264
WOMTL	10.0 14.6	9.2 ± 16.4	11.8 ± 15.2	0.305	10.2 ± 13.5	7.5 ± 9.7	0.343	0.484
WOMKPR	2.6 ± 3.1	2.9 ± 3.1	2.5 ± 3.1	0.415	3.3 ± 3.3	1.3 ± 2.6	0.004	0.022
WOMSTFL	1.2 ± 1.6	1.0 ± 1.5	1.4 ± 1.7	0.170	1.5 ± 1.4	1.1 ± 1.4	0.207	0.321
WOMADLL	7.0 ± 10.7	6.5 ± 12.1	8.2 ± 11.1	0.358	6.9 ± 10.1	5.2 ± 6.6	0.380	0.526

Age at MENR = how old at the time of meniscal surgery, Age at INJ = how old at the time of the injury, INJ-MENR = time difference (years) between time point of the injury and meniscal surgery, MENR-MRI = time difference (years) between time point of the meniscal surgery and MRI scan.

Continues data are expressed as mean \pm SD. Categorical data are presented in numbers of participants with percentage in parentheses. ***p-values were calculated using either chi-square or one-way ANOVA as appropriate. *-**p-values were calculated using either Pearson's χ 2-test or independent t-test as appropriate. Ptcp = Participants. PASE = Physical activity score of the elderly, WOMTL = WOMAC total score, WOMKRP = WOMAC knee pain, WOMSTFL = WOMAC stiffness, WOMADLL = WOMAC disability score.

^{*}p-value = indicates differences between traumatic meniscal surgery and controls with knee trauma.

p-value = indicates differences between degenerative meniscal surgery and controls without knee trauma.

p-value = indicates differences among all groups.

Table 2.

en knees with meniscal surgery related to trauma and menisc

Differences in cartilage T2 values between knees with meniscal surgery related to trauma and meniscal surgery related to degenerative disease

	Traumatic meniscal surgery		Degenerative meniscal surgery		Effect size		
	Adjusted means	[95% CI]	Adjusted means	[95% CI]	Adjusted means	[95% CI]	p-value
Global	32.83	[32.48,33.18]	32.94	[32.42,33.47]	-0.11	[-0.75,0.53]	0.731
Global WB	32.92	[32.55,33.29]	33.23	[32.69,33.78]	-0.31	[-0.98,0.35]	0.354
LF	34.71	[34.20,35.22]	35.15	[34.39,35.91]	-0.44	[-1.37,0.49]	0.351
LT	28.06	[27.59,28.54]	28.81	[28.10,29.51]	-0.74	[-1.60,0.11]	0.088
MF	38.38	[37.85,38.90]	38.03	[37.26,38.80]	0.34	[-0.59,1.28]	0.472
MT	30.48	[29.96,31.00]	30.56	[29.79,31.33]	-0.08	[-1.02,0.86]	0.867
PAT	32.39	[31.73,33.05]	31.82	[30.84,32.81]	0.57	[-0.62, 1.76]	0.346

Data are given as adjusted means, corrected for age, sex, and baseline BMI with [95% confidence intervals]. P-values <0.05 are in bold characters. WB = weight bearing (mean of both femoral and tibial compartments), PAT = patella, MT = medial tibia, LT = lateral tibia, MF = medial femur, LF = lateral femur.

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Table 3.

Differences in cartilage T2 values between participants with meniscal surgery and non-surgical controls

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	Traumatic meniscal surgery		Non surgery controls with preceding trauma		Difference in adjusted means		p-value
	Adjusted means	[95% CI]	Adjusted means	[95% CI]	Adjusted means	[95% CI]	
Global	32.83	[32.48,33.18]	31.89	[31.53,32.24]	0.94	[0.45,1.44]	<0.001
Global WB	32.92	[32.55,33.29]	31.96	[31.59,32.32]	0.97	[0.45,1.48]	<0.001
LF	34.71	[34.20,35.22]	34.22	[33.70,34.74]	0.49	[-0.23,1.22]	0.182
LT	28.06	[27.59,28.54]	27.31	[26.84,27.78]	0.76	[0.09,1.42]	0.025
MF	38.38	[37.85,38.90]	37.25	[36.72,37.77]	1.13	[0.39,1.87]	0.003
MT	30.48	[29.96,31.00]	29.53	[29.01,30.05]	0.95	[0.22,1.68]	0.011
			21.54	[30.86,32.22]	0.85	[-0.09,1.79]	0.076
PAT	32.39	[31.73,33.05]	Non surgery controls	. , .			0.070
PAT	32.39 Degenerative men		Non surgery controls trau	s without preceding	Difference in adj		0.070
PAT			Non surgery controls	s without preceding			0.070
PAT	Degenerative me	niscal surgery	Non surgery controls trau	s without preceding ma	Difference in adj	justed means	<0.001
Global	Degenerative men	niscal surgery [95% CI]	Non surgery controls traus	s without preceding ma	Difference in adj	justed means [95% CI]	
Global Global WB	Degenerative men Adjusted means 32.94	[95% CI]	Non surgery controls traus Adjusted means	s without preceding ma [95% CI] [31.02,32.07]	Difference in adj Adjusted means	[95% CI]	<0.001
Global Global WB LF	Degenerative men Adjusted means 32.94 33.23	[95% CI] [32.42,33.47] [32.69,33.78]	Non surgery controls traus Adjusted means 31.55 31.67	[95% CI] [31.02,32.07] [31.12,32.22]	Adjusted means 1.35 1.56	[95% CI] [0.60,2.09] [0.79,2.33]	<0.001 <0.001
Global Global WB LF LT	Degenerative men Adjusted means 32.94 33.23 35.15	[95% CI] [32.42,33.47] [32.69,33.78] [34.39,35.91]	Non surgery controls traus Adjusted means 31.55 31.67 33.48	[95% CI] [31.02,32.07] [31.12,32.22] [32.70,34.26]	Adjusted means 1.35 1.56 1.67	[95% CI] [0.60,2.09] [0.79,2.33] [0.59,2.75]	<0.001 <0.001 0.003
Global	Degenerative men Adjusted means 32.94 33.23 35.15 28.81	[95% CI] [32.42,33.47] [32.69,33.78] [34.39,35.91] [28.10,29.51]	Non surgery controls traus Adjusted means 31.55 31.67 33.48 27.23	[95% CI] [31.02,32.07] [31.12,32.22] [32.70,34.26] [26.53,27.92]	Adjusted means 1.35 1.56 1.67 1.58	[95% CI] [0.60,2.09] [0.79,2.33] [0.59,2.75] [0.60,2.56]	<0.001 <0.001 0.003 0.053

Data are given as adjusted means, corrected for age, sex, and baseline BMI with [95% confidence intervals]. P-values <0.05 are in bold characters. WB = weight bearing, PAT = patella, MT = medial tibia, LT = lateral tibia, MF = medial femur, LF = lateral femur.