Imaging of articular cartilage

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Abstract

We tried to review the role of magnetic resonance imaging (MRI) in understanding microscopic and morphologic structure of the articular cartilage. The optimal protocols and available spin-echo sequences in present day practice are reviewed in context of common pathologies of articular cartilage. The future trends of articular cartilage imaging have been discussed with their appropriateness. In diarthrodial joints of the body, articular cartilage is functionally very important. It is frequently exposed to trauma, degeneration, and repetitive wear and tear. MRI has played a vital role in evaluation of articular cartilage. With the availability of advanced repair surgeries for cartilage lesions, there has been an increased demand for improved cartilage imaging techniques. Recent advances in imaging strategies for native and postoperative articular cartilage open up an entirely new approach in management of cartilage-related pathologies.

Key words: Cartilage; chondromalacia; magnetic resonance imaging

Introduction

With an increase in availability of treatment options for the articular cartilage lesions, imaging takes the front seat in determining those lesions. There has been a constant advancement in field of articular cartilage imaging.

Morphology

The articular surfaces of joints in human body are lined by hyaline cartilage. The entire role of articular cartilage is to distribute force, reduce friction, and prevent wear. It is a connective tissue of very specialized nature and configuration.[1]

Molecular Basis

Hyaline cartilage is composed of water, which constitutes about 80%, and remainder is cellular substance, intercellular matrix, and ground substance, which are essentially responsible to give firmness to cartilage. The absence of lymphatics, nerve endings and blood vessels does not allow its regeneration. When a joint is compressed, water is redistributed through cartilage cushion to absorb the compressive force whilst the overall cartilage volume may remain unchanged.[2]

There are multiple layers in articular cartilage. The superficial layer has lower proteoglycan concentration and has no permeability for water. The superficial zone may act as a cushion distributing the impact of compressive forces. If the forces remain for a prolonged period, then there is a propensity of the impact to be distributed to the deeper zones.[3]

The radial zone is the largest of all and is the deepest in location. The collagen fibers are perpendicular in orientation in upper two-thirds, whereas they are curved obliquely in the lower third. The “tidemark” junction is a thin calcified layer deep to the radial zone. In this layer, the collagen fibers anchor the underlying subchondral bone.[3]

MRI Appearance of Normal Articular Cartilage

It is an immense challenge to image the thin articular cartilage, which has the maximum thickness not beyond 4 mm with smooth curved contours. Magnetic resonance
imaging (MRI) has the privilege of being the radiological investigation with maximum promise in the evaluation of articular cartilage. The merits of MRI are its high spatial resolution and a superb soft tissue contrast, together with multiplanar facility. The sensitivity of MRI is reduced due to the partial volume averaging effects, which may often mask early chondral lesions like fissures and cartilage flaps. To ascertain the efficacy of MRI to produce high-quality images, it is very essential to optimize the spatial resolution for an appropriate signal-to-noise ratio (SNR). The availability of high field strength magnets together with dedicated surface coils and dedicated pulse sequences has taken articular cartilage imaging to a higher level.

The difference in water content between the superficial and deep layers has been thought to be responsible for the homogenous appearance of articular cartilage on MRI [Figure 1A and B].

**Conventional Pulse Sequences**

Common MRI sequences for cartilage constitute T1-weighted, proton density, and T2-weighted imaging sequences, with or without fat saturation, but the preferable information is expected to be better assessed on high-resolution imaging which is considered important for morphological and physiologic information. In present day practice, combination of morphologic imaging sequence with high-resolution facility compatible to evaluate the matrix is preferred and is readily available.

2D Fast Spin Echo

To assess the cartilage integrity, it is preferred to have proton density and T2-weighted sequences, with and without fat saturation.4,5 Both intermediate and T2-weighted fast spin-echo (SE) imaging sequences, with and without fat suppression have been advocated in the assessment of articular cartilage integrity.6,7 [Figure 3].

The disruption of ultrastructure of collagen matrix together with intra-articular water imbibement is reflected as areas of increased signal, with morphologic contour defects most likely reflective of increased intra-articular free water and collagenous ultrastructure disruption.

There is a limited role of T1-weighted imaging in cartilage evaluation due to lack of good tissue contrast from the adjacent joint fluid and the articular cartilage surface, making it suboptimal for particular usage [Figure 4]. Additional shortcomings are its limitations in assessing the adjacent soft tissue structures.8

Proton density-weighted sequence has the feature of T1 and efficacy of T2, based upon the local concentration of the hydrogen nuclei [Figure 5]. The intact articular cartilage has low signal intensity as compared to the bright fluid signal intensity, increasing the conspicuity of the focal surface lesions7,9 [Figure 6].

The T2-weighted imaging sequences show excellent contrast difference due to fluid-cartilage interphase, but at the expense of slightly reduced signal from the articular cartilage. T1-weighted conventional SE sequences have high SNR and good spatial resolution, but poor contrast between cartilage and joint fluid.10

**Fat Suppression Techniques**

To improve the dynamic range of signal intensity from an image, fat suppression sequence is implemented in the routine protocol for cartilage imaging pulse sequences. There is better detection of minor signal intensity modifications.11,12 A T1-weighted inversion recovery sequence, termed as short tau inversion recovery (STIR), is another technique by which the effect of fat suppression is obtained. Combination of fat suppression together with three-dimensional (3D) spoiled gradient echo sequence only depicts the articular cartilage as a bright structure.
allowing a better quantitative measurement of cartilage abnormalities and reducing partial volume artifacts.[14]

**Volumetric Sequences**

Optimal spatial resolution of images can be obtained by acquiring thinner slices (0.2 to 0.4 mm thickness) producing isotropic voxels with equal dimensions in the three planes. The advantage of volumetric image acquisition is that it facilitates multiplanar reformation without loss of the spatial resolution.

**Gradient Echo Sequences**

Gradient echo sequences allow volumetric image acquisition with reduced imaging times and improved spatial resolution. Technically, a refocusing pulse separates two or more gradient echoes, eventually combining the echoes to generate an image.[15,16] Gradient-recalled echo (GRE) sequences suffer from a potential fallacy, as they are unusually sensitive and susceptible to intravoxel dephasing, especially in patients with previous surgical intervention or some hardware placement.[13]

**3D Fast Spin Echo Imaging**

The structures that are oblique in course or some small structures that extend into multiple slices pose a great amount of problem. To tackle this problem, there is an advent of 3D sequences which are fast SE images equipped with flip angle modulation reducing blurring and imaging times, making isotropic imaging possible. Such 3D images have an isotropic resolution and an advantage of multiplanar reformatting.

**Figure 2 (A and B):** Axial FS T2W images of the patella targeted for cartilage imaging (A) large FOV (B) small FOV. Note better delineation of chondral morphology in retropatellar cartilage when images obtained at a smaller FOV. Similarly, use of a small FOV for superficial structures like joints may help us improve spatial resolution and conspicuity of chondral based lesions.

**Figure 3:** Sagittal FS T2W FSE image of knee joint. The articular cartilage appears hyperintense. Note excellent morphological detail of articular cartilage. The other structures of knee joint can also be identified. However, finer morphology of articular cartilage along tidemark junction (arrow) is compromised.

**Figure 4:** Coronal T1W image of knee joint. The articular cartilage appears hypointense. The bone details like bone trabecular pattern (blue arrow), marginal osteophytosis (white arrow) and subchondral sclerosis (yellow arrow) are nicely depicted. However, there is compromised information of articular cartilage.
SPGR

Spoiled gradient echo sequences such as SPGR or fast low angle shot (FLASH) provide T1-weighted images with better contrast between cartilage (hyperintense) and intra-articular fluid (hypointense).\[^{19,20}\] Standard SE MRI has a lower accuracy as compared to the fat-suppressed SPGR. The sensitivity of the sequence approaches 93%, with articular cartilage appearing bright and remainder of the structures appearing relatively dark\[^{21,22}\] [Figure 8]. 3D SPGR is recommended by the International Cartilage Repair Society (ICRS) as a standard sequence in imaging for evaluation of the articular cartilage lesions, especially in post cartilage repair status. The images are primarily T1-weighted sequences obtained at a low flip angle. The disadvantages are longer acquisition times and inadequacy to depict surface defects as well as the other joint structures. Finally, 3D gradient echo methods are less useful for the diagnosis of ligament or meniscal tears than SE techniques. Despite these limitations, 3D SPGR imaging is considered the standard for morphologic imaging of cartilage\[^{23}\] [Figure 9].

MR Arthrography

Post cartilage repair, MR arthrography (MRA) is recommended in evaluation of cartilage integrity. The instilled contrast outlines cartilage defects and their contours. A small amount of 2mmol/l gadolinium contrast (Dotarem, Guerbet LLC, USA) mixed with saline is injected within the joint space to increase its sensitivity to detect cartilage abnormalities\[^{24,25}\]. MRA has been shown to be more sensitive than the standard 3D T2-weighted gradient echo sequences for the detection of cartilage lesions\[^{26}\] [Figure 10].

Pathomechanics

Chondral lesions in their early stages result in contour changes in the form of swelling, fibrillation, and surface irregularity, together with cartilage thinning.

Degenerative Disease

Chronic stress-related microtrauma to articular cartilage and bone results in arthritis and osteoarthritis, which involve diffuse chondropathy, osteophytes, eburnation of the bone surface, and subchondral changes\[^{28}\] [Figure 11].

Trauma

Microtrauma-related articular fractures and thinning of articular cartilage is the common etiopathogenesis of osteoarthritis. Chondral fractures are well demonstrated at arthroscopy in patients with hemarthrosis\[^{28}\].
On MRI, chondral pathologies are depicted as areas of focal chondral defects or deeper fractures involving the subchondral bone [Table 1]. Chondral lesions are graded based upon the thickness of articular cartilage disruption. The cartilage may undergo swelling (focal or diffuse) represented as an area of hyperintensity on fluid-sensitive sequence [Figure 12]. Focal disruption of cartilage may be a linear cleft or multiple linear clefts representing fissures of variable depths [Figure 13]. Full-thickness cartilage separation from the tidemark zone results in chondral flap injuries or delamination of articular cartilage [Figure 14]. Associated with injury to articular cartilage, there may often be injury to subchondral bone, which may be represented as bone marrow edema, subchondral bone fracture, or displaced fracture [Figure 15].

Osteochondral lesions need a further detailed description based upon location, size, and depth of lesion [Figure 16A and B].

**New Pulse Sequences**

To date, no one acquisition sequence has been proven to be ideal, although the majority of publications on volume and thickness measurements have used the 3D T1-weighted SPGR sequence.[29,30]

These methods include the following:
- Steady-state free precession (SSFP) [fast imaging employing steady-state acquisition (FIESTA)]
- True fast imaging with SSFP (true-FISP)
- Balanced fast field echo and its variant, fluctuating equilibrium MRI (FEMR)
Multi-echo techniques such as dual excitation in the steady state [dual-echo steady state (DESS)]; driven equilibrium techniques such as driven equilibrium Fourier transform (DEFT) and fast recovery fast spin echo; echo-planar techniques such as 3D echo-planar imaging with fat suppression and 3D DEFT; and 3D fast SE methods.[33]

**Diffusion MRI Techniques**

The movement of water within extracellular matrix (ECM) of articular cartilage has been considered a potentially useful measure to assess biochemical constitution of the articular cartilage.

**DWI**

There is alteration of the magnetic field gradients due to the movement of the water. Eventually, this water movement has been calculated as a b-factor, which has an echo time characterizing the degree of T2 weighting and is mapped pixel by pixel.[32,33] Diffusion-weighted imaging (DWI) has been demonstrated as a potential method for assessing cartilage degeneration in vivo[34] and monitoring its repair following surgery.[35]

**ADC**

Apparent diffusion coefficient (ADC) determines the interaction between water molecules in relation to the adjacent environment. This coefficient has been measured in articular cartilage, suggesting integrity of tissue morphology. An increased diffusivity suggests structural disintegration of the ECM and, hence, can be readily utilized as a potential method in post cartilage repair imaging[36,37] with increased diffusivity linked to structural degradation of the ECM.

**DTI**

Orientation of the collagen fibrils in the ECM of the articular cartilage has been studied using diffusion tensor imaging (DTI). The disruption of the collagen network is mapped on DTI. But its drawbacks are intricate data analysis and a longer duration of image acquisition.[38] DTI has been used to study the effects of compression on the cartilage collagen network.[39]

**DESS**

DESS imaging has proven useful for the evaluation of cartilage imaging and 3-mm slices can be acquired in about 6 min. Initial studies of cartilage morphology have been performed using DEFT imaging,[40,41] but this technique has not been conclusively proven to be superior to 2D approaches [Figure 17].

**Balanced SSFP Imaging**

It is an efficient 3D MR sequence obtained with a high signal and fast acquisition times, employing a precision of steady state [Figure 18]. A higher resolution is obtained by multiple acquisitions at an expense of increased scanning time.[42]

Several studies have shown the utility of the balanced SSFP (bSSFP) sequence for imaging articular cartilage.[43-45] Because of the bright synovial fluid and 3D nature of the acquisition, bSSFP may also be useful for imaging internal...
derangements of other structures including ligaments and menisci.\[45\]

**Relaxation Time Mapping**

Water molecules excite and relax reaching back to the equilibrium state. T1 and T2 relaxation times describe the exponential time constants obtained due to movement of the water molecules in a tissue at the field strength.

**T1 Relaxation**

During the spin, the rate at which the nuclei diffuse the energy to the surroundings following the excitation is determined by T1 relaxation, which is sensitive to the macromolecular structure of the cartilage matrix.\[47,48\]

**T2 Relaxation**

The function of the water and collagen content of the

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Figure 11 (A-D): Multiplanar FS PD-weighted images in a 56 year old woman who previously underwent arthroscopic partial medial meniscectomy. MRI images of knee joint show established osteoarthritis in medial-lateral femorotibial and patellofemoral articulation. There is a near-complete loss of articular cartilage covering weight bearing surfaces of femoral and tibial condyles as well as non-weight bearing surface of patellofemoral articulation exposing subchondral bone. There are established marginal osteophytosis and subchondral bone sclerosis

Figure 12: Axial image acquired with multiple-echo data image combination sequence shows grade 1 cartilage lesion. There is a focal area of chondral softening revealed as an area of focal hyperintensity (arrow) within the articular cartilage

Figure 13: Axial 2D fat-saturated PD-weighted fast spin-echo image of knee joint in a 47 year old man shows arthroscopically confirmed superficial cartilage fibrillation (arrows) categorized as grade 2 cartilage lesion

Figure 14: Coronal FS PD weighted image of the knee reveals a grade 2 chondromalacia of the weight-bearing surface of the medial femoral condyle. (arrows) Note the fibrillar contour of cartilage surface in a 36 year old man with surgically confirmed superficial partial-thickness cartilage lesion
The T2 relaxation values are variable, based upon the articular cartilage thickness comparable to the distribution of the water and proteoglycans in the ECM. The measured distribution of the T2 relaxation is depicted as areas of increased or decreased content of the water directly proportional to the degree of the cartilage disintegration. It is of utmost importance to select the MRI technique that has high accuracy to measure T2 relaxation.

**T1rho Imaging**

T1rho imaging is a potentially sensitive MRI technique to detect early depletion of the proteoglycans in the ECM. It detects the alterations in the relaxation of the spins in a magnetic field. This technique is different from T2 mapping that determines the changes in the collagen architecture; moreover, the T1rho value variations are remarkably larger in cases of osteoarthritis, offering a wider spectrum of disease detection.

**Physiologic MRI of Cartilage**

Physiologic MRI helps to explore the internal architecture of the articular cartilage by the use of collagen and proteoglycan arrangement. There is a combination of 70% water and the remainder consisting of type 2 collagen fibers and proteoglycans. The concept of positively charged sodium ions from the collagen fibers interacting with the negatively charged gadolinium contrast particle indirectly depicts the concentration of the proteoglycans.

The interaction is determined by the magnetization transfer and can be readily identified under 1.5 and 3 T systems.

**Delayed Gadolinium-enhanced MRI of the Cartilage**

Early degeneration of articular cartilage occurs in the form of degradation of proteoglycans in ECM. There is a direct relationship of the contrast-enhanced T1 and glycosaminoglycan (GAG) concentration affected by patient’s body mass index. There is an effective penetration of contrast into area of cartilage, which has a disrupted concentration of GAG. A normal articular cartilage has a low gadolinium diethylenetriaminepentaacetic acid (GdDTPA) concentration, whereas areas of damaged articular cartilage allow contrast material to trickle in areas of delayed hyperenhancement of articular cartilage. Imaging-wise, a double dose of the Gd contrast material is injected intravenously and images are acquired after a delay of about 2 h, allowing sufficient time for contrast to percolate into areas of damaged collagen. The concentration of GAG is inversely proportional to concentration of Gd contrast on T1W imaging. A T1 map is plotted based on contrast concentration indirectly quantifying GAG in cartilage. Delayed gadolinium-enhanced MRI of cartilage (dGEMRIC) has a significant promise to assess biochemical changes in articular cartilage morphology and may reflect early cartilage degeneration or maturation based upon concentration of GAG.

**Sodium MRI**

Articular cartilage has a high concentration of sodium. Taking advantage of this, sodium MRI has been brought in for cartilage imaging. The areas of proteoglycan depletion are represented as areas of high sodium concentration. This}

Figure 15: Axial fast spin-echo PD FS image in 39 year old man reveals a wide area of full-thickness chondromalacia (arrow) involving the median ridge and the medial patellar facet categorized as grade 3 lesion (fissuring of cartilage to level of subchondral bone) in the posterior aspect of the patella. The subchondral bone is not exposed, which discriminates this lesion from a grade 4 lesion.
technique is considered an alternative method to dGEMRIC to assess focal cortical defect. Triple-quantum-filtered imaging of sodium in cartilage, which may be even more sensitive to early changes than sodium imaging, is also possible.[57]

3 T versus 1.5 T

SNR is directly proportional to magnetic field strength [Figure 20]. A greater spatial resolution with equal imaging time and SNR is obtained with higher magnetic strengths.[58]

Higher field strength magnets have a potential for an improved articular cartilage imaging. Lower field

Figure 17: Sagittal water selective 3D DESS imaging of wrist (TR/TE 14.1/5 ms). At the flip angle of 60°, there is highest signal intensity of articular cartilage with highest contrast-to-noise ratio of cartilage. Note excellent details of articular cartilage. Note that the details of subchondral bone are compromised.

Figure 18: 47 year old male volunteer with knee pain. Sagittal balanced steady-state free precession (bSSFP) image of knee joint (0.3 mm isotropic resolution and 8 minute scanning time) shows superficial partial-thickness cartilage thinning (arrow) on anterior weight bearing surface of lateral femoral condyle.

Figure 19 (A-C): Axial T1 TSE image (600/6.6, NEX 2.0, 180° flip angle) of the retropatellar cartilage obtained in a volunteer reveals morphologically normal cartilage (A). Corresponding T2 maps for the early (B) and late unloading (C). Note no visible differences over time.

Figure 20 (A and B): Axial 2D FS PD TSE images obtained at a 3T (A) and 1.5T (B) respectively. Note the excellent contrast differentiation among cartilage surfaces, synovial fluid, and subchondral bone, as well as variation of signal intensity within the cartilage. It is very obvious that high field 3T magnet (a) depicts the finer details of the articular cartilage to a greater extent as opposed to a magnet of lesser strength. If all parameters remain similar, the voxel size in a 1.5 T remains 276% larger than in 3.
strength magnets reduce the benefits, especially in non-fat-suppressed methods. With higher field magnets, there is an increase in T1 relaxation times together with increase in frequency differences amongst cartilage, water and marrow fat. There is a twofold increase in SNR at the same spatial resolution and imaging time or a reduction by fourfold in the imaging time.

**Discussion**

MRI non-invasively provides significant information regarding contents of articular cartilage and its macromolecular structure. Availability of cartilage-specific sequence allows sensitive prediction of the mechanical integrity of the articular cartilage.99

Morphologic assessment of articular cartilage by MRI specific sequences provides an excellent tissue contrast that gives detailed information of early chondral-based lesions or pathologies. An appropriate cartilage-based sequence should provide details of cartilage morphology, cartilage thickness, volume, and any disturbances of internal signal intensities. Many of the MRI parameters can assess macromolecular detail, biochemical configuration, and the physiological changes of cartilage.100 In addition to morphologic assessment, tissue parameters that can be measured by MRI techniques also have the potential to provide biochemical and physiologic information about cartilage.

Physiological MRI imaging sequences are available for hyaline cartilage, which readily give information about composition of articular cartilage, collagen matrix, and GAGs. On the other hand, conventional MRI sequences do not help much in cartilage assessment as they typically lack required spatial resolution or information about cartilage physiology.

Hyaline cartilage is made up of chondrocytes and water, together with macromolecules such as glycoproteins, proteoglycans, and collagen. The various layers within articular cartilage are histologically categorized as superficial, transitional, and deep zone, based upon variations in orientations of collagen fibers and number of chondrocytes.

Current musculoskeletal MRI protocols include multiple planes of 2D FSE and 3D gradient echo images. In this article, current understanding of MRI appearance of articular cartilage and its relationship to microscopic and macroscopic structure of articular cartilage, the optimal pulse sequences to be used, and appearance of both degenerative and traumatic chondral lesions are discussed. Improvements have been made in morphologic imaging of cartilage in terms of contrast, resolution, and acquisition time. These improvements allow detailed maps of cartilage surface to be developed that can be used to quantify both thickness and volume. The choice of a particular protocol for imaging articular cartilage depends greatly on patient factors. Currently available MRI imaging techniques for evaluating morphologic and compositional characteristics of articular cartilage in knee in clinical and research settings are also reviewed in the article.

Disorders may involve either the cartilage (chondral disorder), subchondral bone (subchondral bony disorder), or both (osteochondral disorder). Injury of bone is often associated with cartilage damage and vice versa.

Recent MRI techniques include GRE sequences, cartilage mapping, and dGEMRIC sequences. Use of contrast medium either injected intravenously or within the joint is showing promise in evaluation of cartilage damage and GAG glycoprotein content.100

MRI is the imaging method of choice in evaluation of cartilage lesions and alterations in their morphologic composition.

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**Figure 21 (A-E):** MRI images depicting grading of articular cartilage lesions based upon Modified Outerbridge classification. Grade 0 (21A, normal articular cartilage), grade 1 (21B, chondral surface fibrilation), grade 2 (21C, partial thickness chondral lesion), grade 3 (21D, full thickness cartilage lesion reaching the bone), grade 4 (21E, full thickness cartilage lesion exposing the subchondral bone)
It is a very useful investigation in cartilage-based pathologies as well as in follow-up to assess treatment response. Compositional assessment techniques include T2 mapping, delayed gadolinium-enhanced MR imaging of cartilage (dGEMRIC), T1 imaging, sodium imaging, and DWI.

There are various modifications in MRI protocols and parameters which have been formulated to alter the SNR amongst structures surrounding articular cartilage in relation to cartilage proper, indirectly allowing us to ascertain subtle cartilage-based lesions [Figure 21]. MRA of knee helps to increase sensitivity for cartilage lesions and for detection of intra-articular bodies and their donor sites, but it is also not routinely performed. Indications for MRA include evaluation of articular cartilage.

Conclusion

There are several variations in the existing imaging techniques with alterations in the imaging parameters. Numerous variations of these imaging techniques exist, with the imaging parameters and acquisition sequences varying between centers on the basis of multiple factors, including user preferences and the potential strengths and weaknesses of the available vendor hardware platforms. In general, practical MRI protocols for the evaluation of hyaline articular cartilage include at least one cartilage-sensitive sequence obtained in the sagittal plane. Supplemental acquisitions or reformating of 3D data sets in the axial and coronal planes may help to optimize evaluation of patellar cartilage and central weight-bearing aspects of the femoral condyles and tibial plateau, respectively.

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