CARTILAGE IMAGING (S TRATTNIG, SECTION EDITOR)

UTE MRI of the Osteochondral Junction

Won C. Bae · Reni Biswas · Karen Chen · Eric Y. Chang · Christine B. Chung

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Abstract The osteochondral junction is composed of numerous tissue components and serves important functions relating to structural stability and proper nutrition in joints such as the knee and spine. Conventional MR techniques have been inadequate at imaging the tissues of the osteochondral junction primarily because of the intrinsically short T2 nature of these tissues, rendering them "invisible" with the standard acquisitions. Ultrashort time to echo (UTE) MR techniques acquire sufficient MR signal of osteochondral tissues, thereby allowing direct evaluation. This article reviews the anatomy of the osteochondral junction of the knee and the spine, technical aspects of UTE MRI, and the application of UTE MRI for evaluation of the osteochondral junction.

Keywords Magnetic resonance imaging · Musculoskeletal · Knee · Cartilage · Spine · Endplate

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W. C. Bae (⋈) · R. Biswas · K. Chen ·

E. Y. Chang · C. B. Chung

Department of Radiology, University of California, San Diego, 408 Dickinson St., San Diego, CA 92103-8226, USA

e-mail: wbae@ucsd.edu

R. Biswas

e-mail: biswas.reni@gmail.com

K. Chen

e-mail: karenchanchen@gmail.com

E. Y. Chang

e-mail: ericchangmd@gmail.com

C. B. Chung

e-mail: cbchung@ucsd.edu

W. C. Bae·K. Chen·E. Y. Chang·C. B. Chung VA San Diego Healthcare System, 3350 La Jolla Village Drive MC-114, San Diego, CA 92161, USA

Introduction: Osteochondral Junction

The osteochondral junction predominates in many joints in the human body where cartilage attaches to the subchondral bone. These include diarthrodial joints such as the knee and the hip as well as fibrocartilaginous joints such as the intervertebral discs. The anatomy of the osteochondral junction is complex, as several tissue components exist near the junction. For example, in a vertical cross section of the femoral condyle, layers of tissues (Fig. 1a) including uncalcified cartilage (which has superficial, middle and deep layers [1]), calcified cartilage and subchondral bone are present. Similarly, in the spine, layers of tissues (Fig. 1b) including the disc proper (nucleus pulposus and/ or annulus fibrosus), uncalcified and calcified cartilaginous endplates, and subchondral bone of vertebral endplate exist. The term "osteochondral junction" is rather loosely defined and may refer to one or more tissue components in the region.

In the knee, the osteochondral junction (Fig. 1a) serves to connect the overlying articular cartilage to the subchondral bone via calcified cartilage. The deepest layer of softer uncalcified cartilage has radially oriented collagen fibrils [1] that embed into much stiffer calcified cartilage at the tidemark or the calcification front. Calcified cartilage interdigitates with the subchondral bone, and the rough interface between the tissues combined with high stiffness [2] of the calcified cartilage provides mechanical stability and resistance against shear failure [3]. Calcified cartilage is a thin (100-200 µm thick [4•]) tissue, is metabolically active [5], consists of a large proportion of hydroxyapatite $(\sim 65\% [6])$, and remodels with loading [7] and aging [8]. Lane et al. [4•] has showed that calcified cartilage thickness decreases with age, from $\sim 200 \, \mu m$ at about age 20, down to less than 100 µm by age 90. Subchondral bone consists



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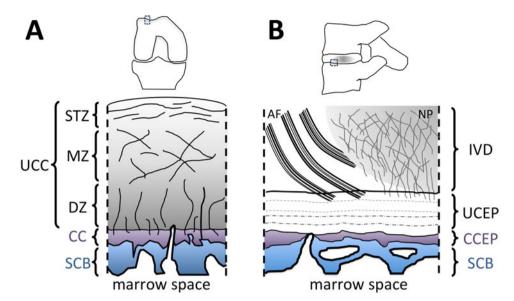


Fig. 1 Anatomy. a Osteochondral junction of the knee. Uncalcified articular cartilage (UCC) consists of three layers: the superficial tangential zone (STZ), middle zone (MZ) and deep zone (DZ). The UCC is attached to the subchondral bone (SCB) via calcified cartilage (CC), which interdigitates with subchondral bone and forms a tight junction. There are vascular canals penetrating the SCB and CC, which allow diffusion of solutes between the deep zone cartilage and

the marrow space. **b** Osteochondral junction of the spine or discovertebral junction. Components of the intervertebral disc (IVD) including the annulus fibrosus (AF) and nucleus pulposus (NP) anchor into the uncalcified cartilaginous endplate (UCEP), which is attached to the calcified cartilaginous endplate (CCEP), supported by the subchondral bone (SCB)

mostly of impermeable cortical bone and provides underlying support for uncalcified and calcified cartilage. The thickness of the subchondral bone plate is on the order of 2 mm, but varies with location, age and joint degeneration [9]. Vascular canals penetrate both the subchondral bone and calcified cartilage, providing a nutritional pathway between the bone marrow and articular cartilage.

The osteochondral junction of the knee has been implicated in the pathogenesis of osteoarthritis. In addition to age-related changes in calcified cartilage [4•], joint loading can induce injury or degenerative changes at the osteochondral junction, including subarticular fracture [10], cloning of the cell in the calcified cartilage layer [11] and cell death in the deep layer [12]. It is also important for cartilage repair. During microfracture surgery in animals, the presence of the calcified cartilage layer is an important determinant of the outcome of the surgery [13].

A similar structure is also seen at the disco-vertebral junction (Fig. 1b) of the spine. Here, the nucleus pulposus and annulus fibrosus anchor into the uncalcified cartilaginous endplate (CEP), which is attached to the calcified CEP, supported by the subchondral bone. The uncalcified CEP is ~ 1 mm thick, and its extracellular matrix consists mainly of proteoglycans, collagen and water. Calcified CEP is $\sim 100~\mu m$ thick and interdigitates with the subchondral bone, similar to the calcified cartilage of the knee. The detailed collagen network architecture of this region has been documented by Inoue et al. [14•] and Wade et al. [15•].

Due to avascular nature, the intervertebral discs must receive a large proportion of nutrition via disco-vertebral junction. A rich supply of blood vessels exists [16], penetrating the subchondral bone and the calcified CEP. Studies have shown a correlation between the low density of vascular canals with disc degeneration [17], emphasizing a nutritional role of the disco-vertebral junction and implication for disc degeneration. Other age-related changes in the CEP have been suggested, such as thinning and occlusion of vascular canals by ectopic calcification [18]. A non-invasive and sensitive method to detect changes in the osteochondral junctions of the human joints would be of considerable value for early diagnosis and understanding of the pathogenesis.

Limitation of Conventional MRI and Advantage of UTE MRI

While the importance of the osteochondral junction for proper function and degeneration of human joints has been recognized, it has been difficult to evaluate this important region using conventional magnetic resonance imaging (MRI) techniques. Tissues of the human body have intrinsic MR properties, such as T1 (spin–lattice relaxation time constant), T2 (spin–spin relaxation time constant) and T1rho (spin–lattice relaxation time constant in rotating frame) [19]. MR pulse sequences, by adjusting parameters



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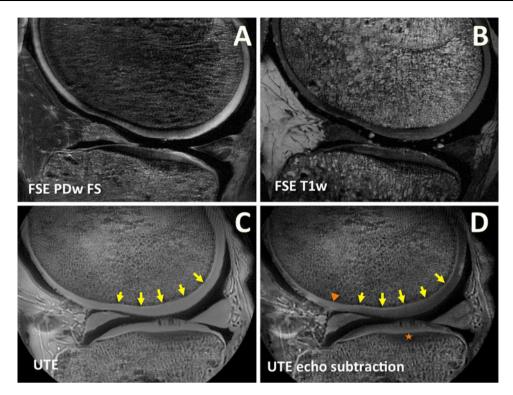


Fig. 2 MR images of a human knee in the sagittal plane obtained using conventional and UTE techniques. Conventional fast spin echo (FSE) **a** proton density-weighted fat suppressed (PDw FS; TR = 2,000 ms, TE = 30 ms) and **b** T1-weighted (T1w; TR = 500 ms, TE = 15 ms) MR images are often used for clinical evaluation. These conventional MR images exhibit low signal intensity at the osteochondral junction and are more susceptible to magic angle effects. **c** UTE MR image obtained at TR = 300 ms and TE = 0.008 ms exhibit many soft tissues with high signal intensity,

including the osteochondral junction (*arrows*). In UTE images, the magic angle artifact in the articular cartilage is reduced compared to conventional MR images. **d** After subtracting the second echo image (at TE = 8 ms) from the UTE image, the osteochondral junction is seen with much greater contrast due to suppression of longer T2 signals. In the subtraction image, osteochondral regions of abnormal morphology can be readily identified, including a region with signal loss (*triangle*) and another region with diffuse thickening (*star*)

of time to repeat (TR), time to echo (TE) and time to spin lock (TSL), are able to obtain images depicting many structures of the joint with preferred contrast. For example, Fig. 2a and b shows images of a knee, showing articular cartilage with different signal intensities relative to the surrounding tissues.

However, due to technical limitations of clinical MR scanners, many tissues have remained "invisible" (i.e., have very low signal intensity) on conventional MR images. A primary reason for this is the intrinsic T2 or T2* of the tissue being too short to provide a detectable signal when imaged with a conventional sequence with relatively long TE (e.g., Fig. 2a, b). In the knee, the bulk of articular cartilage has intermediate T2 values of ~ 50 ms (Fig. 3a), while the layer of calcified cartilage has T2 value ~ 2 ms [20]. Similarly, the nucleus pulposus of the intervertebral disc has a long T2 value ~ 150 ms, while both uncalcified and calcified CEPs have much shorter T2 values, less than 5 ms (Fig. 6). Subchondral bone has even shorter T2 values than 1 ms [21]. Given that a practical minimum TE is ~ 10 ms for a spin echo sequence and ~ 2 ms for a

gradient echo sequence, many short T2 tissues near the osteochondral junction will be "invisible" or imaged with suboptimal contrast because of the rapidly decaying MR signal.

Ultrashort TE (UTE) techniques allow acquisition of MR signals much earlier after excitation than with conventional sequences. With UTE sequences, the minimum TE can now be as short as 8 μs [21]. As a result, MR signal from previously "invisible" tissues with very short T2 values can now be acquired and imaged with high signal intensity (Figs. 2c, d and 4c, d). The majority of tissues near the osteochondral junction, including the deepest layer of the articular cartilage, calcified cartilage, and uncalcified and calcified CEP, can now be directly imaged using UTE techniques.

UTE MR Techniques

A number of UTE techniques have been developed and refined, focusing on the method of image acquisition,



Fig. 3 Quantitative MRI of the knee. Conventional a spin echo multi-echo T2 map (SE; TR = 2,000 ms,TE = 10-80 ms, 8 TEs) and **b** spiral chopped magnetization preparation (SCMP; TR = 1,500 ms, TE = 3 ms,TSL = 10-40 ms, 4 TSLsshow relatively high T2 and T1rho values of articular cartilage. In comparison, quantitative c UTE T2* (TR = 100 ms,TE = 0.012-60 ms, 16 TEs)and d UTE T1rho (TR = 100 ms. TE = 0.012.TSL = 0.02-10 ms, 4 TSLsare able to detect short T2 components in articular cartilage and other soft tissues

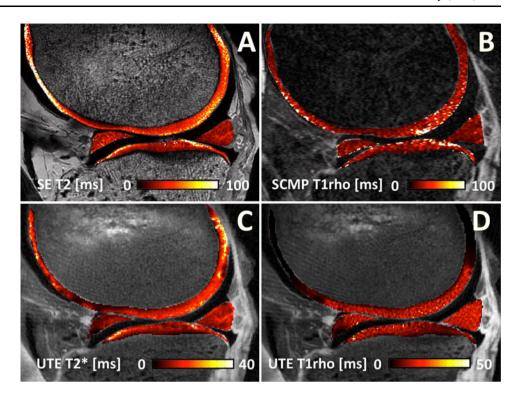
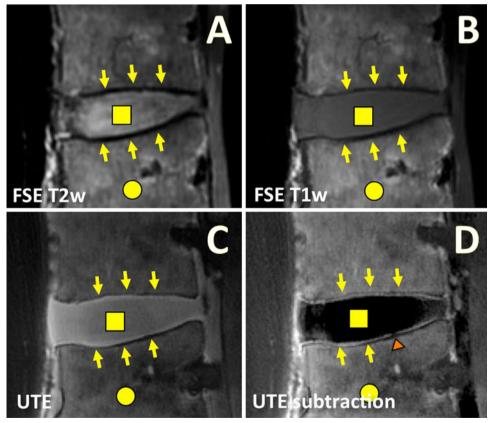


Fig. 4 Morphologic MRI of a human lumbar spine in the sagittal plane using conventional and UTE techniques. Conventional fast spin echo (FSE) a T2-weighted (T2w; TR = 2,000 ms,TE = 80 ms) and **b** T1weighted (T1w; TR = 600 ms, TE = 10 ms) MR images are often used for clinical evaluation of the disc proper (square) and the bone marrow (circle). These conventional MR images exhibit low signal intensity at the disco-vertebral junction (arrows). c UTE MR images obtained at TR = 300 ms and TE = 0.01 ms reveal high signal intensity at the discovertebral junction (arrows) as well as the disc proper (square). d After subtracting the second echo image (at TE = 11 ms) from the UTE image, the osteochondral junction is seen with a much greater contrast. A focal region of abnormal discovertebral junction (triangle) can



contrast enhancement, and quantitation of MR properties. Methods of image acquisition include two-dimensional (2D) [22–24] and three-dimensional (3D) [25, 26]

approaches. The 2D UTE sequence typically employs half excitation pulses with radial sampling of k-space from the center out [27, 28]. Data from the two half excitations are



now be seen

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Table 1 Morphologic UTE MRI for evaluation of human musculoskeletal tissues

Organ or tissue	Technique (Scan parameters)	Findings	Ref.
Knee	3D UTE-weighted subtraction (TR = 6 ms, TE = 0.07 and 3.8 ms)	Determined weighting factors optimized for cartilage, tendon, bone, meniscus	[35]
Mandibular condyle	3D UTE (TR = 50 ms, UTE = 0.05 ms, FOV = 4 cm, matrix = $384 \times 384 \times 384$)	Condylar fibrocartilage imaged with high signal intensity, allowing determination of 3D contour	[78]
Disco-vertebral junction	2D UTE subtraction (TR = 300 ms , TE = 0.008 and 6.6 ms)	Identified uncalcified and calcified cartilaginous endplates as sources of high linear signal intensity	[74••]
Lumbar spine	3D UTE (TR = 4.8 ms , TE = 0.14 ms)	Found association between the presence of CEP defects and disc degeneration	[77]
Tibial cortical bone	3D UTE: subtraction (TR = 20 ms, TE = 0.06 and 4.6 ms), dual-band saturation (TR = 240 ms), inversion recovery (TR = 300 ms, TI = 100 ms)	Subtraction and saturation techniques provided good short T2 SNR and CNR. Inversion recovery had the lowest SNR but uniform short T2 contrast	[79]
Knee	2D UTE Spiral subtraction (Cor, $TR = 100 \text{ ms}$, $TE = 0.36 \text{ ms}$, 10 ms)	In healthy subjects, deep layer of articular cartilage was visualized	[80]
Meniscus	3D UTE subtraction (TR = 35 ms, TE = 0.056 , 5.1 ms)	Meniscal calcifications detected with high signal intensity	[61]
Knee	3D AWSOS UTE (TR = 80 ms , TE = 0.6 – 0.8 ms , inplane voxel = 0.14 mm)	Feasibility of high resolution UTE imaging using AWSOS sequence	[81]
Lower leg, knee, Achilles tendon, forearm	3D UTE subtraction, rescaled subtraction, long T2 water and fat saturation, single inversion recovery, double inversion recovery	Feasibility shown for in vitro and in vivo imaging of osteochondral junction, aponeuroses, meniscus, tendon, ligament and cortical bone	[25]
Tooth	3D SWIFT (TR = 2.5 ms , TE = 0.006 ms)	Hard and soft tissues of teeth were simultaneously imaged	[82]
Wrist (cortical bone)	WASPI	WASPI technique allows imaging of solid bone matrix and tendons, with suppression of soft tissues	[83]
Achilles tendon and cortical bone	2D UTE OSC and subtraction (TR = $200-300$ ms, TE = 0.008 ms)	UTE-OSC technique suppresses long T2 water and fat effectively, creating high contrast for short T2 tissues	[84]
Foot, ankle, knee	3D UTE subtraction (TR = 7.7 ms, TE = 0.05 ms, 2.3 ms at 3T), 3D UTE magnetization preparation (long T2 and fat suppression)	Both techniques well suited for long T2 suppression	[32]
Tibial cortical bone and ankle	2D UTE with inversion recovery pulse (TR = 300 ms, TE = 80 $\mu s)$	T2 suppression allows good visualization of bone and tendon	[38]
Achilles	2D UTE subtraction (TR = 500 ms; TE = 0.08, 6.0, 11.1, 17.7 ms)	UTE MRI shows pathological changes and not seen in conventional MRI	[63]
Spine	2D UTE subtraction (TR = 500 ms, TE = 0.08, 6.0, 11.1, 17.7 ms)	UTE sequence shows hyper-intense band near cartilaginous endplates	[85]
Meniscus	2D UTE subtraction (TR = 500 ms, TE = 0.08, 6.0, 11.1, 17.7 ms), with and without contrast administration	Increase in SI of red zone and perimeniscal tissue after contrast injection, especially at shorter TEs	[86]

2D Two dimensional, 3D three dimensional, UTE ultrashort time to echo, IR inversion recovery, DIR dual inversion recovery, TR time to repeat, TE time to echo, TI time to invert, FOV field of view, SNR signal-to-noise ratio, CNR contrast-to-noise ratio, CEP cartilaginous endplate, AWSOS acquisition weighted stack of spirals, SWIFT sweep imaging with Fourier transformation, WASPI water- and fat-suppressed proton projection MRI

subsequently combined to produce a single radial line of k-space. The radial k-space data are converted to a Cartesian grid and reconstructed by 2D inverse Fourier transformation (FT). The 3D UTE sequences usually employ a short hard pulse excitation followed by 3D radial ramp sampling [25, 26, 29] or 3D cones trajectory [30], followed by re-gridding of k-space onto a Cartesian grid and 3D inverse FT.

These image acquisition sequences may be used in conjunction with techniques to modulate image contrast. In

particular, suppression of the long T2 signal while preserving the short T2 signal has been of interest in order to better evaluate short T2 tissues. A simple and effective technique is to acquire two images at varying echo times and perform digital image subtraction [31•, 32]. The first echo image (obtained at the minimum TE, typically less than 0.1 ms) contains MR signals from both short and long T2 tissues (Figs. 2c, 4c), while the second echo image (obtained at longer TEs, typically greater than a few ms) contains the signal mainly from the long T2 tissues since



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the signal from the short T2 tissues has mostly decayed away. Subtracting the second image from the first image thus yields an image unmasking short T2 tissues (Figs. 2d, 4d). For this technique, short T2 contrast can be modulated by varying TEs, usually the later TE [33]. A variation of this technique, such as rescaled or weighted digital subtraction [34, 35], is also available. Other advanced techniques involving long T2 water saturation [36, 37] or inversion nulling of water [25, 38] or water and fat using dual adiabatic inversion recovery (DIR) [39] have also been introduced. It should be noted that short T2 tissues often have short T1 values. For example, in the human spine, the T1 value of the nucleus pulposus is $\sim 1,250$ ms, while that of uncalcified CEP is ~ 550 ms [40]. Such T1 differences can also be used to accentuate signal intensity from short T2 tissues by optimizing TR and the flip angle [40, 41]. Table 1 summarizes applications of morphologic UTE MRI for imaging of various musculoskeletal tissues.

UTE techniques also enable MR quantification of short T2 tissues. Quantitative techniques have been developed to determine UTE T2* and UTE T1rho values. T2* is typically determined by exponential fitting of UTE MR images

obtained at different TEs [42, 43]. Similarly, UTE T1rho can be measured through exponential fitting of UTE T1rho images acquired at a series of spin-lock times (TSLs) [44]. Figure 3c and d shows UTE T2* and UTE T1rho maps of a knee slice, respectively. With a sufficient number of images, it is also possible to simultaneously determine short and long T2* components. UTE bicomponent analysis [45] utilizes a model that has a reduced number of fitting parameters and noise correction to determine both the value and fraction of short and long T2* components. An increasing number of studies seek to determine multicomponent T2* properties for musculoskeletal tissues [46, 47]. Table 2 summarizes the application of quantitative UTE MRI for imaging of various musculoskeletal tissues.

UTE MR Evaluation of the Osteochondral Junction of the Knee

Conventional MRI is routinely used to evaluate injury and repair of the articular cartilage of the knee [48–51]. Sequences such as fat-suppressed proton density-weighted

Table 2 Ouantitative UTE MRI for evaluation of musculoskeletal tissues

Organ or tissue	Technique (scan parameters)	Findings	Ref.
Meniscus and cartilage in meniscal repair	2D UTE T2* (TR = 350 ms, TE = 0.3, 6.3, 12.5, 18.7 ms)	Higher UTE T2* values in repaired menisci and articular cartilage.	[60]
Patella: zone of calcified cartilage (ZCC)	2D DIR UTE T2* (TR = 300 ms, TI1 = 140 ms, TI2 = 105 ms, TE = $0.008-12$ ms); 2D DIR UTE T1rho (TSL = $0.02-12$ ms)	Characterization of patellar ZCC UTE T2*: 2.0 ± 1.2 ms UTE T1rho: 3.6 ± 1.2 ms	[87]
Meniscus calcification	2D IR UTE T2* (TR = 300 ms, TE = 0.012, 0.05, 0.1, 0.2, 0.4 ms); 3D UTE T2* (TR = 35 ms, TE = 0.056, 2, 5.1, 7.1 ms)	Lower T2* value of calcification (~ 10 ms) compared to surrounding tissue (15–30 ms)	[61]
Meniscus	3D AWSOS UTE T2* (TR = 30 ms, TE = 0.6–40 ms, 11 TEs)	Meniscal T2* values were higher in ACL-injured subjects, with and without concomitant tear	[62]
Cortical bone (lower extremity)	2D UTE T2* bicomponent analysis for short and long T2* and fractions (TR = 200 ms, TE = $0.008-8$ ms, 16 TEs); 3D UTE for total water content	UTE measures of water content and T2* fraction correlated with porosity and biomechanical measures of cortical bone	[55]
Knee articular cartilage	3D AWSOS UTE T2* (TR = 80 ms, TE = 0.6–40 ms, 11 TEs)	Repeatability of UTE T2* values of cartilage in asymptomatic subjects was determined	[43]
Cortical bone (lower extremity)	3D hybrid-radial UTE T2* (TR = 20 ms, 0.25–1.65 ms, 8 TEs)	Repeatable bone water quantification in vivo	[64]
TMJ disc	2D UTE T2* (TR = 500 ms , TE = $0.1-15 \text{ ms}$, 8 TEs)	Significant correlation between UTE T2* and indentation stiffness of TMJ disc was found	[88]
Achilles	2D UTE T1rho (TR = 400 ms, TE = 0.008 ms, TSL = 0.2, 2, 5, 10 ms)	UTE T1rho ranged from 3.0 ± 0.5 ms for healthy volunteers to 5.2 ± 0.6 ms for specimens	[44]
Meniscus	2D UTE T1rho (TR = 400 ms, TE = 0.008 ms, TSL = 0.2, 2, 5, 10 ms)	UTE T1rho of 7.98 \pm 1.43 ms found in healthy volunteers	[44]
Achilles	2D UTE (TR = 500 ms , TEs = $0.14-2.5 \text{ ms}$, 8 TEs)	UTE T2* values of 2.9 and 3.0 ms were found	[63]

2D two dimensional, 3D three dimensional, UTE ultrashort time to echo, IR inversion recovery, DIR dual inversion recovery, TR time to repeat, TE time to echo, TI time to invert, FOV field of view, AWSOS acquisition-weighted stack of spirals, TMJ temporomandibular joint



(PDw; long TR and short TE) (Fig. 2a) and T1-weighted (T1w; short TR and short TE) (Fig. 2b) spin echo or fast spin echo sequences are used to evaluate morphologic changes in articular cartilage as well as trabecular bone and bone marrow. However, tissues near the osteochondral junction are seen with low signal intensity in these conventional MR images. Additionally, the magic angle effect [52, 53] results in orientation-dependent fluctuations in signal intensity of the overlying cartilage, which further confounds evaluation of the osteochondral junction.

In contrast to conventional MR images, UTE MRI of the articular cartilage enables direct visualization of deep layers. Using the shortest TE of 0.008 ms (Fig. 2c), UTE MRI of a knee exhibits high signal intensity from all layers of the articular cartilage, meniscus, and surrounding connective tissues. In addition, it is apparent that the UTE MR image is less sensitive to the magic angle effect; similar signal intensity of articular cartilage is seen throughout the femoral condyle. In the image, the morphology of the osteochondral junction (Fig. 2c, arrows) is not well defined because of the confounding signal intensity from longer T2 tissues. After digital image subtraction of the second echo image (obtained at TE = 8 ms) to suppress long T2 signals, the resulting image (Fig. 2d) accentuates the signal intensity from short T2 tissues. Specifically, the osteochondral junction is seen with distinct, continuous and high signal intensity (Fig. 2d, arrows). In some samples, focal regions of abnormal morphology can be found, including thinning/absence of the signal intensity (Fig. 2d, triangle) or diffuse thickening (Fig. 2d, star). In a recent study performed on cadaveric patellae [54•], ~80 % of the combined length of all osteochondral junctions had a normal morphology, $\sim 10 \%$ had thinned/absent morphology and the remainder had a thick diffuse morphology.

It remains to be determined how different UTE MR morphologies and properties are related to the structure and health of the osteochondral junction as well as other tissues of the knee. In an early validation study [54•], UTE MR morphology of experimentally prepared samples containing variable tissue components were compared. The following samples were prepared: intact osteochondral (uncalcified cartilage, calcified cartilage, subchondral bone), uncalcified cartilage, calcified cartilage and subchondral bone, and subchondral bone. Only the samples containing the uncalcified cartilage (at the deepest layer) and calcified cartilage exhibited the characteristic high linear signal intensity, suggesting those two tissue components are sources of the signal intensity. It should be noted that while the cortical bone is detectable using UTE techniques [55–57] when imaged on its own, when imaged alongside other short T2 tissues such as calcified cartilage, the subchondral bone exhibits a relatively low signal intensity. In another preliminary study [58], the UTE MR

morphology of the osteochondral junction of the human patella was compared with histological measures of calcified cartilage thickness and roughness. Regions that exhibited the MR appearance of thinning or absence of the linear signal intensity had a thinner calcified cartilage as well as a smoother interface to the subchondral bone.

Quantitatively, UTE MRI has often been performed to evaluate full thickness articular cartilage or other tissues of the knee as a whole, but not specifically to evaluate the osteochondral junction. This is partly due to the thinness of the osteochondral junction of the knee, combined with the long scan duration required for achieving high spatial resolution in quantitative sequences. In a few in vitro studies that have evaluated UTE MR properties of the patellar osteochondral junction, UTE T1 values ranging from 250 to 400 ms, UTE T2* values ranging from 1 to 3 ms and UTE T1rho values ranging from 2 to 5 ms were found in cadaveric patellar samples [20]. When UTE T2* and UTE T1rho values of the osteochondral junctions with normal versus abnormal UTE morphology were compared [59], abnormal regions with diminished signal intensity in subtraction images also had ~40 % greater UTE T2* and T1rho values than the normal regions. Quantitative UTE MR techniques are increasingly used to evaluate other parts of the lower extremities, including the overlying articular cartilage [43, 60], the meniscus [44, 61, 62], the ligaments [44, 63] and the cortical bone [55, 64]. Additional studies are needed to fully understand implications of MR changes of the osteochondral junction for the health of the knee.

UTE MR Evaluation of the Osteochondral Junction of the Spine

Conventional MRI is routinely used to evaluate intervertebral disc degeneration and associated conditions. Based on the MR signal intensity and morphology, disc grading [65, 66] can be performed, usually based on spin echo images in the sagittal plane (Fig. 4a, b). Conditions such as disc herniation and nerve root compression can be detected accurately [67, 68] using similar techniques. Conventional quantitative techniques have also been useful for evaluation of disc degeneration: T2 values of the disc correlate strongly with the water content [69, 70] and proteoglycan content [71, 72]. Similar relations were found using T1rho imaging [73]. While useful for evaluation of the nucleus pulposus and anulus fibrosus, and gross morphology of the spine, conventional sequences yield very little signal from the region of the disco-vertebral junction (Fig. 4a, b, arrows). It should be noted that while it is feasible to image uncalcified CEP using conventional gradient echo imaging at a short TE [40], the image contrast for the CEP may be suboptimal.



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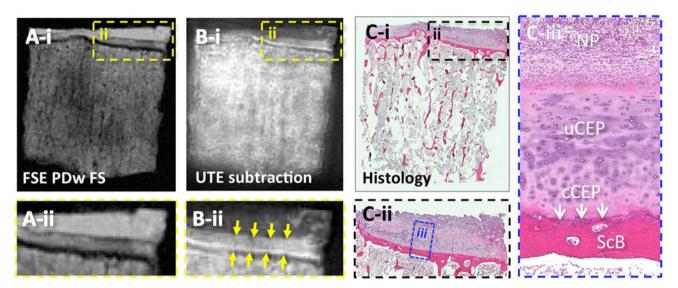


Fig. 5 Comparison of MRI and histology of a small disco-vertebral specimen. **a** Conventional fast spin echo proton density-weighted fat-suppressed (FSE PDw FS; TR = 2,000 ms, TE = 15 ms) image exhibits a medium to low signal intensity at the disco-vertebral junction. In contrast, **b** the UTE subtraction (TR = 300 ms, TE = 0.008 and 6.6 ms) MR image exhibits a medium to high signal intensity with a bilaminar appearance. On the UTE subtraction image, the thicker layer with medium signal intensity (**b-ii**, *downward arrows*) corresponded with the thicker layer of the uncalcified

cartilaginous endplate, and a thinner layer (\mathbf{b} - $\mathbf{i}\mathbf{i}$, $upward\ arrows$) of high signal intensity corresponded with the thinner layer of the calcified cartilaginous endplate. These observations were consistent with \mathbf{c} the histology of the sample, stained with hematoxylin and eosin. In \mathbf{a} through \mathbf{c} , \mathbf{i} is the full-size image, and $\mathbf{i}\mathbf{i}$ are magnified images. NP nucleus pulposus, uCEP uncalcified cartilaginous endplate, cCEP calcified cartilaginous endplate, scb subchondral bone

Ultrashort time-to-echo (UTE) MR techniques have enabled direct imaging of the osteochondral junction of the spine or disco-vertebral junction. Similar to knee imaging, the UTE image of the spine at the shortest TE reveals the entire disc (Fig. 4c, square) as well as the CEP (Fig. 4c, arrows) with medium to high signal intensity. Using the subtraction technique, the longer T2 signals are diminished, and the osteochondral junction is seen with a characteristic continuous, linear, and high signal intensity (Fig. 4d, arrows). With enhanced contrast, focal regions of abnormal morphology are more easily detected (Fig. 4d, triangle), unlike in the first echo UTE image (Fig. 4c). Abnormal morphologies including signal loss (Fig. 4d, triangle), thickening or irregularity have been noted. A histologic validation study [74••] has been performed on small disco-vertebral specimens (Fig. 5) containing the disc, uncalcified and calcified CEPs, and subchondral bone. This study confirmed that the CEPs (uncalcified and calcified) exhibit lowered signal intensity in conventional fast spin echo MRI (Fig. 5a), while the UTE subtraction image revealed two layers of characteristic high signal intensity (Fig. 5b-ii), which corresponded with thicker uncalcified and thinner calcified layers of CEP. In the corresponding histology (Fig. 5c), the structure and arrangement of these tissues at the osteochondral junction can be seen. Using a different combination of prepared samples, the study [74••] determined that both uncalcified and calcified CEP contributed to the characteristic UTE MR morphology, while the disc

tissue and subchondral bone did not. MR quantification of the disco-vertebral junction is also difficult because of the thinness of the region, but has been performed in a limited number of small specimens at a high spatial resolution (Fig. 6). Compared to a conventional multi-echo spin echo T2 sequence that yielded images (Fig. 6a–c) of the disco-vertebral junction with low signal intensity, both UTE T2* (Fig. 6f–h) and UTE T1rho (Fig. 6k–m) images show the region with high-to-medium signal intensity. The T2 map created from a conventional sequence (Fig. 6d) shows highly noisy results in the CEP, while the UTE T2* (Fig. 6i) and UTE T1rho (Fig. 6n) maps show smoother results, likely to be more accurate. Curve fitting (Fig. 6e, j, o) in the CEP region of interest suggested a UTE T2* value of 2.9 ms and a UTE T1rho value of 4.5 ms in the region.

Implications of the altered UTE MR morphology or MR properties of the disco-vertebral junction for spine health have yet to be determined. There is early evidence suggesting that abnormal UTE morphology is related to formation of calcium deposits in the CEP [75], determined by micro CT imaging. Such deposition may lead to occlusion of vascular canals and hindered nutrient transport into the disc. More importantly, in vitro [76] and in vivo [77] UTE MR studies have also shown that a significant association exists between abnormal UTE morphology and advanced degeneration of the adjacent discs. Increased UTE abnormality was also found with more aged subjects. Future UTE MRI studies may offer important insights



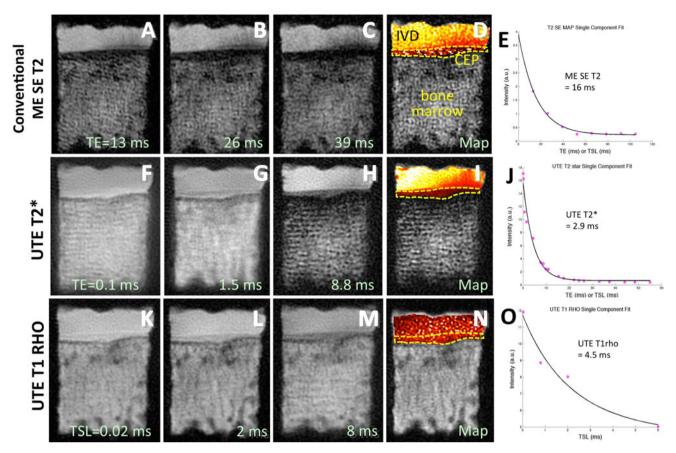


Fig. 6 Quantitative MRI of the disco-vertebral junction. **a–c** Conventional multi-echo spin echo T2 (ME SE T2) images taken at increasing TEs exhibit low signal intensity in the region of the cartilaginous endplate (CEP) (**d**, *dotted line*). **d** The resulting T2 map shows a lot of noise, and **e** the mean T2 value is relatively high at 16 ms. In contrast, **f–h** UTE T2* and **k–m** UTE T1rho images are

able to depict the cartilaginous endplate with markedly higher signal intensity. Corresponding i UTE T2* and n UTE T1rho maps show consistent values throughout the CEP (dotted lines). j The mean UTE T2* and o mean UTE T1rho values of the CEP were 2.9 and 4.5 ms, respectively

into the epidemiology of the pathology of the disco-vertebral junction and how it is related to spine health and back pain.

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Compliance with Ethics Guidelines

Conflict of Interest Won C. Bae, Reni Biswas, Karen Chen, Eric Y. Chang and Christine B. Chung declare that they have no conflicts of interest.

Human and Animal Rights and Informed Consent This article does not contain any studies with human or animal subjects performed by the authors.

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