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Femoral condyle insufficiency fractures: associated clinical and morphological findings and impact on outcome

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Abstract

Objective To determine the characteristics of femoral condyle insufficiency fracture (FCIF) lesions and their relative associations with the risk of clinical progression.

Materials and methods This HIPAA-compliant retrospective study was approved by our Institutional Review Board. Seventy-three patients (age range, 19–95) were included after excluding patients with post-traumatic fractures, bone marrow infarct, osteochondritis dissecans, or underlying tumor. Two board-certified musculoskeletal radiologists classified morphologic findings including lesion diameter, associated bone marrow edema pattern, and associated cartilage/meniscus damage. Electronic medical charts were evaluated for symptoms, risk factors, and longitudinal outcomes, including total knee arthroplasty (TKA). Imaging characteristics were correlated with clinical findings, and comparison of outcome groups was performed using a regression model adjusted for age.

Results The majority of patients with FCIF were women (64.4 %, 47/73), on average 10 years older than men (66.28 ±15.86 years vs. 56.54±10.39 years, p=0.005). The most common location for FCIF was the central weight-bearing surface of the medial femoral condyle; overlying full thickness cartilage loss (75.7 %, 53/70) and ipsilateral meniscal injury (94.1 %, 64/68) were frequently associated. Clinical outcomes were variable, with 23.9 % (11/46) requiring TKA. Cartilage WORMS score, adjacent cartilage loss, and contralateral meniscal injury, in addition to decreased knee range of motion at presentation, were significantly associated with progression to TKA (p<0.05).

Conclusions FCIF are frequently associated with overlying cartilage loss and ipsilateral meniscal injury. The extent of cartilage loss and meniscal damage, in addition to loss of knee range of motion at the time of presentation, are significantly associated with clinical progression.

Keywords Spontaneous osteonecrosis of the knee · Insufficiency fracture · Femoral condyle · Magnetic resonance imaging

Introduction

Spontaneous osteonecrosis of the knee (SONK) was introduced as a distinct entity by Ahlbäck et al. in 1968 [1]. The authors described a subset of elderly patients presenting with acute onset of unprovoked severe knee pain maximally tender over the weight-bearing surface of the medial femoral condyle. Radiographs obtained during the first weeks were normal; however, subsequent flattening of the medial femoral condyle and development of subchondral lucency with a well-circumscribed sclerotic margin appeared after several months. After excluding entities such as osteochondritis dissecans, osteoarthritis, fracture, infection, neuropathy, and tumor, the authors concluded that the condition represented an atypical spontaneous manifestation of femoral osteonecrosis.

Since the introduction of SONK, there has been much inquiry into its pathophysiology. Traumatic and vascular insults are thought to be the most plausible etiologies, although the
absence of a clear precipitating factor suggests a multifactorial pathogenesis [2, 3]. In 1999, Yamamoto et al. described subchondral insufficiency fractures of the femoral head within lesions previously classified as osteonecrosis [4]. This discovery prompted a reevaluation of SONK at the femoral condyle [2, 5, 6]. A spectrum of femoral condylar pathology has since been proposed, with FCIF and osteonecrosis being recognized as distinct entities along this spectrum, and osteoporosis was implicated as a risk factor [2, 7–10]. Currently, there remains limited knowledge about the clinical findings and morphological abnormalities associated with femoral condyle insufficiency fractures (FCIF). In particular, meniscal tears appear to be frequently encountered with these lesions, although existing studies are limited by small numbers and low-field-strength magnetic resonance (MR) imaging systems [7].

Though treatment options vary widely and include oral/intravenous bisphosphonate treatment, high tibial osteotomy, unicompartmental knee arthroplasty, and total knee arthroplasty (TKA), no studies to date have explored the ways in which MR findings of FCIF lesions can be used to predict the longitudinal evolution of these lesions and how they are associated with treatment [11–16]. Our objectives were therefore to (i) analyze MR characteristics of FCIF, (ii) describe associated cartilage, bone marrow, and ligamentous abnormalities, and (iii) correlate these findings with concomitant clinical symptoms, risk factors, and longitudinal outcomes. We hypothesized that elucidation of coexisting imaging and clinical findings would further our understanding of the pathophysiology and prognosis of the disease process, thus helping to guide clinicians towards a more appropriate prognosis-based treatment plan.

Materials and methods

Subjects

The Institutional Review Board approved this HIPAA-compliant retrospective study. The requirement for informed consent was waived. A picture archiving and communications system (PACS) database search from August 2000 through August 2013 was performed for the keywords “femoral condyle insufficiency fracture,” “spontaneous osteonecrosis knee,” and/or “SONK” in the radiology report impression. This search yielded 162 patients; 84 met the typical inclusion criteria of an insufficiency fracture previously described by Yamamoto et al. [2]; these include a low-signal band on T1-weighted imaging adjacent to the subchondral bone in the weight-bearing portion of the femoral condyle associated with significant amount of bone marrow edema pattern (BMEP) with high signal on T2-weighted imaging (Fig. 1). Patients with non-insufficiency type fracture, bone marrow infarct, osteochondritis dissecans (defined as a central osteochondral separation and/or fragmentation), or underlying tumor were excluded [17]. Technically limited MR images (motion artifact, phase encoding artifact, research protocols) were also excluded. This ultimately yielded 73 patients for inclusion (Fig. 2).

Clinical data

All 73 subjects underwent retrospective review of their clinical data documented within the electronic medical record. “Baseline” was defined as the initial orthopedic visit to address knee pain. Pain was described by duration of symptoms and the visual analog pain scale [18]. Historical information including previous trauma, steroid administration (intra-articular versus systemic), and prior knee surgery was recorded. Body mass index (BMI), vitamin D, and dual-energy X-ray absorptiometry (DXA) analysis of bone mineral density were also noted. Records and follow-up radiographs were reviewed to determine healing of FCIF or disease progression. In particular we analyzed the number of patients which went on to require TKA within the follow-up period. Patients in the non-TKA group with less than 6 months of follow-up were excluded from the outcomes analysis.

An orthopedic surgeon or sports medicine physician performed a physical examination of the knee at baseline. Objective indicators of the knee exam were recorded according to the New Knee Society Knee Scoring System, including alignment, instability, range of motion, flexion contracture, and extensor lag [19].

MR imaging

Subjects underwent MR imaging of the knee at 3 Tesla (n=35) and 1.5 Tesla (n=38) (GE Healthcare, Waukesha, WI, USA) in an eight-channel knee coil (In-Vivo, Gainesville, FL, USA). The following sequences were used for image analysis: (1) coronal intermediate-weighted (iw) 2D fast spin-echo (FSE) sequence with fat saturation, (2) coronal T1-weighted 2D FSE sequence, (3) sagittal proton density-weighted 2D FSE sequence, (4) sagittal iw 2D FSE sequence with fat saturation and (5) axial iw 2D FSE sequence with fat saturation (Table 1).

Image analysis

MR images were reviewed on PACS workstations (Agfa, Ridgefield Park, NJ, USA) and assessed for the presence and location of FCIF, meniscal, cartilage, and ligamentous lesions as well as the presence of bone marrow edema pattern (BMEP), synovitis, and other pathology. Two board-certified musculoskeletal radiologists with 6 years of experience analyzed the studies independently. In cases of divergent findings,
consensus readings with a third board-certified musculoskeletal radiologist (23 years of experience) were performed.

FCIF were characterized in terms of morphology, location, and size. Location was described according to a zone system detailed by Ramnath and Kattapuram, which is outlined in Fig. 3 [6]. Involvement of the weight-bearing versus non-weight-bearing condyle surface was noted. Size measurements in two dimensions were obtained from coronal and sagittal fat-saturated FSE images, along with the maximum depth of the defect, measured from the articular surface to the meniscus (asterisk) is consistent with associated meniscal injury. An 84-year-old female with left knee pain after a fall. Coronal T1-weighted image demonstrates markedly increased bone marrow edema pattern (dashed outline) within the medial femoral condyle with fracture line (white arrow). Abnormal intrinsic signal, tear, and extrusion of the medial meniscus is again shown (asterisk).

Fig. 1  
(a) A 67-year-old female with knee pain. AP radiograph demonstrates radiolucent ovoid lesion with sclerotic halo within the medial femoral condyle (white arrow). (b) An 84-year-old female with left knee pain after a fall. Coronal T1-weighted image demonstrates a low signal intensity focus beneath the articular surface of the medial femoral condyle (white arrow). Abnormal intrinsic signal and extrusion of the medial meniscus (asterisk) is consistent with associated meniscal injury. (c) Corresponding fat-suppressed intermediate-weighted image from patient in (b) demonstrates markedly increased bone marrow edema pattern (dashed outline) within the medial femoral condyle with fracture line (white arrow). Abnormal intrinsic signal, tear, and extrusion of the medial meniscus is again shown (asterisk).

Fig. 2  
Flowchart demonstrates patient inclusion and exclusion criteria.

PACS database search from 8/1/2000-8/31/2013 for knee MR reports of “femoral condyle insufficiency fracture”, “spontaneous osteonecrosis knee”, and/or “SONK” (n=162)

Inclusion exams (n=84)
- Femoral condyle insufficiency fracture as demonstrated on MR imaging.

Excluded exams (n=11)
- Non-insufficiency (i.e. post-traumatic) fracture (n=3)
- Bone marrow infarct (n=4)
- Osteochondritis dissecans (n=0)
- Underlying tumor (n=1)
- Technically limited (n=3)

Patients with MR demonstrating spontaneous femoral condyle insufficiency fracture (n=73)

Patients with clinical and/or radiographic follow-up of at least 6 months, or who progressed to TKA prior to 6 months of follow-up (n=46)

Patients in the non-progression group who did not have at least 6 months of clinical or radiographic follow-up (n=27)
most proximal aspect of the fracture line in either coronal or sagittal images.

BMEP, cartilage defects or thinning, ipsilateral meniscal pathology, and ligamentous injury were characterized. BMEP was classified as none, mild (involving less than one-third of the femoral condyle, excluding the trochlea), moderate (one-third to two-thirds of the femoral condyle) or severe (greater than two-thirds of the femoral condyle) [20]. Cartilage loss was described according to the Modified Outerbridge classification system with size further characterized as either involving less than or greater than 50 % of the fracture diameter for overlying cartilage pathology, or involving less than or greater than 1 cm in greatest sagittal dimension for adjacent cartilage pathology [21]. The presence of cartilage fracture (defined as chondral separations involving the full thickness of the articular cartilage down to subchondral bone) or delamination as well as a Whole Organ MR Score (WORMS) cartilage score for the entire knee were also recorded [22, 23]. The WORMS cartilage score was determined by deriving the maximum value cartilage score from any compartment, with the exclusion of the compartment containing the FCIF lesion. Femoral condyle deformity was assessed in initial and follow-up radiographs based on a previously described grading scheme of articular surface bone attrition as none, mild, moderate, or severe [23].

Meniscal morphology was graded using a six-point scale as an expansion of the WORMS classification: 1, normal, 2, intrasubstance degeneration, 3, simple tear, 4, complex tear without deformity, 5, complex tear with deformity, 6, macerated [23]. Maceration was defined as advanced destruction of the meniscus with abnormal signal, deformity, and tear. Associated meniscal extrusion (greater than 3 mm of the medial meniscus or 2 mm of the lateral meniscus) or meniscal root involvement was noted. Ipsilateral and contralateral collateral ligament integrity was described on a four-point scale: 1, normal, 2, intrinsic signal abnormality but normal thickness, 3, thickening and/or incomplete disruption of fibers, and 4, complete tear. Cruciate ligaments were assessed for signal abnormality, thickening, and/or tear.

The presence of synovitis was noted, manifest by joint effusion, diffuse hyperintense signal in the fat pad on T2/proton density/intermediate-weighted fat-suppressed sequences, thickening of the synovial membranes, and intra-articular synovial bands or debris [24, 25]. Muscle atrophy and edema were also documented.

### Table 1 MR sequence parameters

<table>
<thead>
<tr>
<th>MR sequence</th>
<th>TR (ms)</th>
<th>TE (ms)</th>
<th>FOV (cm)</th>
<th>BW (kHz/pixel)</th>
<th>Slice thickness (mm)</th>
<th>Matrix sizeb (pixels)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Coronal intermediate-weighted 2D FSE with fat saturation(^a)</td>
<td>3200–4600</td>
<td>40–55</td>
<td>130–150</td>
<td>81.37–139.49</td>
<td>4</td>
<td>320×224</td>
</tr>
<tr>
<td>Coronal T1-weighted 2D FSE sequence</td>
<td>550–780</td>
<td>7–15</td>
<td>130–150</td>
<td>61.04–244.14</td>
<td>4</td>
<td>384×256</td>
</tr>
<tr>
<td>Sagittal proton density-weighted 2D FSE</td>
<td>760–1800</td>
<td>15–25</td>
<td>130–150</td>
<td>61.04–81.37</td>
<td>3</td>
<td>320×224</td>
</tr>
<tr>
<td>Sagittal intermediate-weighted 2D FSE with fat saturation(^a)</td>
<td>3000–5000</td>
<td>40–60</td>
<td>130–150</td>
<td>81.37–139.49</td>
<td>3</td>
<td>320×224</td>
</tr>
<tr>
<td>Axial intermediate-weighted 2D FSE with fat saturation(^a)</td>
<td>3000–5000</td>
<td>40–60</td>
<td>130–150</td>
<td>81.37–139.49</td>
<td>4</td>
<td>320×192</td>
</tr>
</tbody>
</table>

\(^a\) Frequency-selective fat saturation

\(^b\) Acquisition matrix

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Fig. 3 A 50-year-old female with a 10-day history of right knee pain. a The lateral and medial femoral condyles are divided into an external, central, and medial zone in the coronal plane to demonstrate the zonal classification system, here demonstrating a lesion in the external zone. b Coronal image of the FCIF measuring 3.5 mm in depth; this measurement (yellow bracket) is obtained by measuring the distance from the articular surface of the condyle to the proximal-most aspect of the lesion. c Schematic representation of location classification modality, with relative frequencies in each section represented by color. Light pink=0–5 %; medium pink=5–10 %; bright red=79.5 %
Statistical analysis

Statistical analysis was performed using IBM SPSS® Statistics 20.0 (IBM, Armonk, NY, USA) and Stata/IC Version 13 (StataCorp, College Station, TX, USA). Descriptive statistics were calculated for each parameter per group, and comparisons of outcome subgroups between predictor variables were performed using a logistic regression model adjusted for age and gender. Statistical significance was defined as $p<0.05$, and a statistical trend as $p<0.10$ [26].

Results

Subject characteristics

Baseline characteristics of the study population are summarized in Table 2. A total of 73 subjects, 26 men and 47 women, were included. Mean patient age was 62.81±14.84 years, and mean BMI was 27.74±5.73. Of note, the female subjects with FCIF were older than the males (mean age 66.28±15.86 compared to 56.54±10.39, $p$ value <0.005).

Sixteen of the subjects (two men and 14 women) underwent DXA to assess bone mineral density. Of these, 13 (81.3 %) demonstrated an abnormal DXA result indicating either osteopenia or osteoporosis based on T-scores [27]. The serum 25-hydroxy vitamin D levels were available for 23 patients, with an average concentration of 24.17±14.38 ng/ml (lab normal reference range 30.0-74.0 ng/ml). Of the patients for whom clinical data on intra-articular and systemic steroid use was available, 21.3 % (10/47) had received at least one prior intra-articular steroid injection into the affected knee, and 6.5 % (3/46) were prescribed ongoing oral steroid therapy.

Table 2  Demographic and clinical characteristics of the subjects

<table>
<thead>
<tr>
<th>Parameter</th>
<th>All patients ($n=73$)</th>
<th>Non-TKA** ($n=35$)</th>
<th>TKA** ($n=11$)</th>
<th>$p$ value$^b$</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gender*</td>
<td></td>
<td></td>
<td></td>
<td>0.086</td>
</tr>
<tr>
<td>Male</td>
<td>35.6 (26/73)</td>
<td>40.0 (14/35)</td>
<td>9.1 (1/11)</td>
<td></td>
</tr>
<tr>
<td>Female</td>
<td>64.4 (47/73)</td>
<td>60.0 (21/35)</td>
<td>90.9 (10/11)</td>
<td></td>
</tr>
<tr>
<td>Age (years)$^c$</td>
<td>62.81±14.84</td>
<td>61.17±14.74</td>
<td>67.91±14.05</td>
<td>0.189</td>
</tr>
<tr>
<td>Body mass index (kg/m$^2$)$^c$</td>
<td>27.74±5.73</td>
<td>28.48±6.66</td>
<td>29.72±3.82</td>
<td>0.290</td>
</tr>
<tr>
<td>Abnormal DXA scan*</td>
<td>81.3 (13/16)</td>
<td>45.5 (5/11)</td>
<td>25.0 (1/4)</td>
<td>0.415</td>
</tr>
<tr>
<td>Joint tenderness present*</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Medial</td>
<td>74.5 (35/47)</td>
<td>78.3 (18/23)</td>
<td>75.0 (6/8)</td>
<td>0.908</td>
</tr>
<tr>
<td>Lateral</td>
<td>23.4 (11/47)</td>
<td>17.4 (4/23)</td>
<td>37.5 (3/8)</td>
<td>0.362</td>
</tr>
<tr>
<td>Both</td>
<td>2.1 (1/47)</td>
<td>13.0 (3/23)</td>
<td>25.0 (2/8)</td>
<td></td>
</tr>
<tr>
<td>Visual analog pain$^a$</td>
<td>5.8±1.8</td>
<td>4.2±3.0</td>
<td>4.7±3.0</td>
<td>0.483</td>
</tr>
<tr>
<td>Prior ipsilateral knee surgery*</td>
<td>16.7 (8/48)</td>
<td>8.0 (2/25)</td>
<td>16.7 (1/6)</td>
<td>0.555</td>
</tr>
<tr>
<td>Prior steroid treatment*</td>
<td>27.7 (13/47)</td>
<td>20.8 (5/24)</td>
<td>0 (0/7)</td>
<td>0.905</td>
</tr>
<tr>
<td>Intra-articular</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Oral</td>
<td>6.4 (3/47)</td>
<td>10.5 (2/23)</td>
<td>0 (0/7)</td>
<td>0.897</td>
</tr>
<tr>
<td>Vitamin D level (ng/ml)$^d$</td>
<td>18.91±16.24</td>
<td>21.38±16.95</td>
<td>13.50±13.44</td>
<td>0.533</td>
</tr>
<tr>
<td>Physical therapy$^a$</td>
<td>65.9 (27/41)</td>
<td>69.6 (16/23)</td>
<td>20.0 (1/5)</td>
<td>0.070</td>
</tr>
<tr>
<td>Range of motion at presentation$^a$</td>
<td>122.8±15.3</td>
<td>123.5±15.8</td>
<td>109.0±12.9</td>
<td>0.038</td>
</tr>
<tr>
<td>Duration symptoms to 1st visit (days)$^d$</td>
<td>228.6±407.2</td>
<td>415.0±818.0</td>
<td>1741.9±1765.3</td>
<td>0.053</td>
</tr>
</tbody>
</table>

DXA dual X-ray absorptiometry, TKA total knee arthroplasty

*Values reported as percentage, with number values in parentheses

**Note that in only 46 patients follow-up of more than 6 months or clinical documentation of TKA was available

$^a$Values reported as mean±standard deviation

$^b$All associations between parameters and outcome (TKA vs. non-TKA) were calculated using a logistic regression model adjusted for age and gender, with the exception of both age and gender which were entered into univariate regression models. Significance defined as $p<0.05$
The medial femoral condyle was the most common site of insufficiency fracture (65/73, 89.0 %), with the majority of cases involving the central, weight-bearing condylar segment (63/73, 86.3 %) (Figs. 1 and 3). The average size of the insufficiency fracture line measured 12.8±5.0 mm in the coronal plane by 18.8±9.1 mm in the sagittal plane, with an average depth of 3.9±2.3 mm.

BMEP, one of the criteria used to define the presence of insufficiency fracture, was present in all cases to varying degrees. Mild edema was present in 40.8 % (29/71) of cases, moderate edema was present in 28.2 % (20/71) of cases, and severe edema was present in 31.0 % (22/71) of cases.

Cartilage loss was strongly associated with the presence of insufficiency fractures, as overlying full-thickness cartilage loss was present in the majority of cases (75.7 %, 53/70). Additionally, half of the cases demonstrated full-thickness loss of cartilage along the affected condyle adjacent to the insufficiency fracture. The overall maximum cartilage WORMS score for the knee (excluding the compartment with the insufficiency fracture) was 4.57±1.36, indicating that the majority of patients had associated large full-thickness cartilage defects in compartments separate from the site of the lesion.

Ipsilateral meniscal pathology was also commonly associated with insufficiency fracture at the knee. Nearly all of the cases (94.1 %, 64/68) demonstrated ipsilateral meniscal pathology on MRI, with over half (54.4 %, 37/68) revealing complete meniscal maceration. By contrast, the contralateral meniscus was much less affected, with 44.3 % (31/70) of cases appearing normal and an additional 28.6 % (20/70) with only mild intrasubstance degeneration.

**Imaging in relation to clinical outcomes**

A summary of the clinical and imaging outcomes is provided in Tables 2 and 3, respectively. While the orthopedists’ clinical notes suggested overall favorable outcomes, with only 4.3 % (2/46) of patients reporting worsened symptoms at time of follow-up, subsequent imaging studies were less optimistic.

<table>
<thead>
<tr>
<th>Table 3</th>
<th>MRI characteristics of femoral condyle insufficiency fractures</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Parameter</strong></td>
<td><strong>All patients (n=73)</strong></td>
</tr>
<tr>
<td>Location of femoral condyle</td>
<td></td>
</tr>
<tr>
<td>Medial</td>
<td>89.0 (65/73)</td>
</tr>
<tr>
<td>Lateral</td>
<td>11.0 (8/73)</td>
</tr>
<tr>
<td>Location of FCIF within condyle</td>
<td></td>
</tr>
<tr>
<td>External</td>
<td>9.6 (7/73)</td>
</tr>
<tr>
<td>Central</td>
<td>86.3 (63/73)</td>
</tr>
<tr>
<td>Mesial</td>
<td>4.1 (3/73)</td>
</tr>
<tr>
<td>Size of fracture band (mm)</td>
<td></td>
</tr>
<tr>
<td>Coronal diameter</td>
<td>12.82±5.02</td>
</tr>
<tr>
<td>Sagittal diameter</td>
<td>18.83±9.11</td>
</tr>
<tr>
<td>Depth</td>
<td>3.90±2.27</td>
</tr>
<tr>
<td>Cartilage quality score</td>
<td></td>
</tr>
<tr>
<td>Overlying cartilage loss</td>
<td>4.33±1.35</td>
</tr>
<tr>
<td>Overlying cartilage defect size (mm)</td>
<td>2.73±0.72</td>
</tr>
<tr>
<td>Adjacent cartilage loss</td>
<td>3.90±1.36</td>
</tr>
<tr>
<td>WORMS maximum cartilage score</td>
<td>4.57±1.36</td>
</tr>
<tr>
<td>Meniscus quality score</td>
<td></td>
</tr>
<tr>
<td>Ipsilateral meniscus</td>
<td>4.57±1.84</td>
</tr>
<tr>
<td>Contralateral meniscus</td>
<td>2.23±1.56</td>
</tr>
<tr>
<td>Bone marrow edema pattern</td>
<td>2.90±0.85</td>
</tr>
<tr>
<td>Presence of muscle edema</td>
<td>16.7 (12/72)</td>
</tr>
<tr>
<td>Presence of joint effusion</td>
<td>89.0 (65/73)</td>
</tr>
</tbody>
</table>

**FCIF** femoral condyle insufficiency fracture; **TKA** total knee arthroplasty; **WORMS** Whole-Organ Magnetic Resonance Imaging Score

*Values reported as percentages, with number values in parentheses

**Note that in only 46 patients follow-up of more than 6 months or TKA was available

*Values reported as mean±standard deviation

*All analyses performed with logistic regression model adjusted for age and gender to determine association of parameter with outcome group (TKA vs. non-TKA). Significance defined as p<0.05

*Calculated as worst cartilage lesion score from unaffected compartment
Follow-up radiographs were available for 46 of the 73 cases, and demonstrated overall unchanged (30.4%, 14/46) or worsened (39.1%, 18/46) deformity of the femoral condyle suggesting progression of the underlying process (Fig. 4). Interestingly, only two of the cases (4.3%) demonstrated interval healing with resolution of the initial insufficiency fracture (Fig. 5).

A total of 46 patients were included in the outcome analysis of this study after excluding 27 patients who did not have at least 6 months of clinical or radiographic follow-up. For patients included in the outcomes analysis, the average follow-up was 36.1±30.1 months; this included an average of 39.9±32.6 months in the non-progression group and 22.5±11.7 months in the progression group. Ultimately, TKA was performed in an additional 23.9% (11/46) of these cases. Initial MR imaging findings that were associated with progression to TKA included adjacent and overall cartilage lesions, as well as meniscal tears. Adjacent cartilage defects were significantly more severe in patients who progressed to TKA (adjacent cartilage loss score 3.6±1.4 in the non-TKA group vs. 4.8±0.4 in the TKA group, \( p = 0.027 \)) and overall WORMS cartilage scores were higher in the TKA group (4.4±1.3 in the non-TKA group vs. 5.7±0.5 in the TKA group, \( p = 0.042 \)). Interestingly, integrity of the contralateral meniscus was also found to be significantly associated with progression as well, with significantly higher meniscal injury scores in the TKA group (1.85±1.26 vs. 3.80±1.81, \( p = 0.039 \)). No significant associations were found with regards to muscle atrophy and edema and risk of FCIF progression.

Clinical variables that were shown to be significantly related to progression of FCIF to TKA included decreased range of motion at time of presentation (123.5±15.8 degrees in the non-TKA group vs. 109.0±12.9 degrees in the TKA group, \( p = 0.038 \)). Physical therapy treatment also trended towards significance, with the non-TKA group being more likely to have sought treatment by a physical therapist (69.6 vs. 20.0%, \( p = 0.070 \)). Association of gender with progression trended towards significance, with females tending to have a higher rate of progression (47.6 vs. 6.7%, \( p = 0.086 \)). Other clinical characteristics, such as presence of abnormal DXA, serum vitamin D level, and visual analog pain scale (VAS), were found to have no significant association with risk of progression to TKA.

**Discussion**

With this retrospective review of 73 cases, we present to the best of our knowledge the largest case series of MRI-diagnosed FCIF reported to date. In addition to demonstrating the frequent association of FCIF with ipsilateral meniscal pathology and full-thickness cartilage loss, our study demonstrated the frequency of FCIF progression to TKA, and the relative infrequency of demonstrable radiographic healing. Patients with signs of diffuse intra-articular knee pathology, in addition to patients with decreased range of knee motion at presentation, were found to have significantly increased risk of progression to TKA.

To understand the demographics of FCIF, it is worth noting that the majority of patients in our case series were female (64.4%) and on average were approximately 10 years older than the male patients. Imaging pathology was typically manifest at the central weight-bearing surface of the medial femoral condyle. The high frequency of osteopenia or osteoporosis among our study population, especially in comparison to previously reported rates of osteopenia or osteoporosis in normal adult populations, supports that the impaired bone architecture along the load-bearing surfaces is likely a risk factor for insufficiency fracture [28, 29]. However, the degree of abnormality (osteopenia versus osteoporosis) to predispose to fracture remains controversial [10, 30]. Additionally, in our study, female gender did show a modest association with progression of FCIF to TKA. The basis for this finding is unclear but may be related to differences in bone quality between male and female subgroups, which has previously been demonstrated in normal adult populations [31, 32].

![Fig. 4](image-url) Natural history of medial femoral insufficiency fracture. Initial intermediate-weighted coronal image in a 50-year-old woman with knee pain (a) demonstrates subchondral insufficiency fracture of the posterior aspect of the medial femoral condyle (arrow). Three months later, follow-up MR (b) demonstrates disease progression with additional site of fracture (asterisk)
FCIF were also strongly associated with ipsilateral meniscal pathology, as nearly all of the cases (94.1%) demonstrated ipsilateral meniscal pathology on MRI, and over half (54.4%) had complete meniscal maceration. Similarly, in our study full thickness cartilage loss was also strongly associated with FCIF lesions, and was found in over 75% of our cohort. These associations support the hypothesis that altered biomechanical forces at the femoral condyle due to a damaged meniscus and cartilage loss can contribute to the development of an insufficiency fracture.

Interestingly, in our outcomes analysis, diffuse cartilage and meniscal health was significantly associated with risk of progression of FCIF. In particular, adjacent cartilage loss and cartilage WORMS score were significantly higher among the progression group, with overlying cartilage loss also showing a mild association with progression as well. The intimate ties between cartilage health and FCIF have previously been described [6]. Though our study does not establish causality in this relationship between cartilage integrity and lesion progression, it does raise the possibility that FCIF lesions may give rise to more rapid cartilage destruction, as is sometimes seen with similar lesions of the femoral head [33, 34].

Knee functionality was also found to be significantly correlated with tendency for FCIF progression. Specifically, increased knee range of motion at the time of presentation was found to be significantly associated with decreased rate of FCIF progression, and attendance of at least one physical therapy session was similarly associated with a lower progression rate. Though again lacking in the ability to determine causality, our study still raises an interesting question as to whether targeted physical therapy sessions designed to increase knee mobility might ultimately prevent or slow FCIF progression, especially when started early during the course of the disease.

Finally, it is interesting to note that characteristics that we anticipated would significantly influence progression rates of FCIF such as lesion depth or diameter, or BMEP extent (which itself can suggest a diagnosis of FCIF), were not found to have a significant relationship to prognosis [35]. This could potentially have important implications for how best to interpret these imaging studies, and reinforces the concept that, in cases of FCIF, the integrity of the surrounding cartilage and intra-articular structures has greater prognostic value than characteristics such as fracture diameter and depth. Of note, with only 46 patients included in our outcomes analysis, our study was not sufficiently powered to detect small differences in these variables between progression and non-progression groups, and thus the potential prognostic value of these variables with regards to FCIF lesions remains unclear. Future multi-center studies enrolling greater numbers of patients may be able to provide a more definitive answer as to whether these imaging characteristics have any association with progression risk.

Our study had several limitations. For one, this study was limited by its retrospective design and relatively small sample size, yet to the best of our knowledge this is the largest study that has been published on FCIF lesions. Another inherent limitation in the retrospective design of this study is the lack of consistency among the clinical follow-up protocol. Because of this lack of a standard protocol, patients had varying degrees of clinical and radiographic follow-up after the initial imaging diagnosis. Additionally, there was no established treatment protocol for modalities such as physical therapy, and patients who attended one physical therapy session could potentially be grouped in the same manner as those who attended multiple sessions on a regular basis.

Another limitation posed by the retrospective nature of the study is our inability to include a normal control cohort for comparison purposes in determining relative frequency of certain imaging findings. However, the frequencies reported in our study differ quite notably from those that have been reported in healthy adult cohorts in previously published studies. For one, our reported frequency of osteoporosis/osteopenia (81.3%) is notably different than that of normal asymptomatic adult populations of similar age ranges, which have previously been reported at 25.8-35% [28, 29]. Similarly, the rate of BMEP in our study population (100%, with 31% having severe BMEP) is notably higher than a
previously published study reporting the frequency of any BMEP in knee MR exams performed in healthy normal adults as 36% [36]. The previously described rates of cartilage loss and meniscal lesions in normal control populations, reported as 1.0-5.3 and 40%, respectively, was also significantly less than the frequencies of these lesions in our study cohort (75.7 and 94.1%, respectively) [36]. We feel that these differences in frequencies effectively demonstrate the relative prevalence of certain imaging findings among FCIF patients.

Finally, though TKA is in itself a verifiable outcome measure, it is not ideal in the sense that the decision to proceed with TKA is based on many factors, some of which are independent of lesion severity. Nevertheless, we feel that this outcome measure generally speaks to the extent of lesion progression in a majority of FCIF patients. Future studies emphasizing a standardized clinical and radiographic follow-up protocol, in addition to a larger sample size, may provide further insight into the prognostic value of FCIF imaging characteristics.

In summary, FCIF are frequently associated with concurrent meniscal and cartilage damage, and coexistence of these abnormalities, along with knee mobility at time of presentation, may be important prognostic indicators of more rapidly progressive lesions. The findings of this study emphasize the importance of close interpretation of MR imaging when dealing with potential FCIF lesions, to establish a proper diagnosis and to consider potential clinical outcomes.

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Conflict of interest The authors declare that they have no conflict of interest.

References


