Introduction
Osteoarthritis is a degenerative condition that increases in incidence and prevalence with age and is a major cause of morbidity in people older than 65 years. This condition most commonly occurs at the knee, particularly in the medial tibiofemoral compartment, and varus knee osteoarthritis accounts for more than 95% of knee osteoarthritis in Japan. Despite the occurrence of significant morbidity, the etiology and factors involved in the pathogenesis of osteoarthritis and the development of varus knee osteoarthritis have not been fully elucidated. There have been few reports that compared cartilage, ligament, and meniscus degeneration and radiographic alignment in patients with severe varus knee osteoarthritis.

The aim of this study was to examine the relationship between cartilage, ligament, and meniscus degeneration and radiographic alignment in severe varus knee osteoarthritis in order to understand the development of varus knee osteoarthritis. The inclusion criteria for the study were primary osteoarthritis in a varus knee and total knee arthroplasty in our hospital. No previous history of anterior cruciate ligament (ACL) damage was found in these patients, and no previous operations had been done in their affected knee joints. There were 6 men and 47 women, with 40 right knees and 31 left knees studied; their mean age at operation was 73.5 years. The ligament, meniscus, degeneration of joint cartilage, and radiographic alignments were examined visually.

Materials and Methods
Over a 4-year period, 53 patients (71 knees) were selected for this study. The inclusion criteria for the study were primary osteoarthritis in a varus knee and total knee arthroplasty in our hospital. No previous history of anterior cruciate ligament (ACL) damage was found in these patients, and no previous operations had been done in their affected knee joints. There were 6 men and 47 women, with 40 right knees and 31 left knees studied. The mean age at operation was 73.5 years (range 60-86 years). Written, informed consent was obtained from all subjects. All the
following data were gathered in these cases. The abbreviations we used in this study are shown in Table 1.

During surgery, we assessed the ligaments, meniscus and cartilage by gross examination. The ACL and PCL were also inspected and divided into 3 groups: intact, ruptured, or disappeared (Table 2). The lateral meniscus (LM) and medial meniscus (MM) were also inspected and divided into 3 groups: intact, degenerated, and ruptured or disappeared.

Degeneration of joint cartilage in the lateral and medial femorotibial joints were also examined visually. Using the International Cartilage Repair Society (ICRS) classification, cartilage degeneration was divided into 5 grades: grade 0 was normal; grade 1 was superficial lesions that included soft indentation and/or superficial fissures and cracks; grade 2 was lesions extending down to less than 50% of cartilage depth; grade 3 was cartilage defects extending down more than 50% of cartilage depth, as well as down to calcified layer and down to but through the subchondral bone; and grade 4 was exposure of subchondral bone.

The femorotibial joint was divided into 4 zones: lateral femoral condyle (LFC), lateral tibial plateau (LTP), medial femoral condyle (MFC), and medial tibial plateau (MTP). Cartilage degeneration was examined in each area. Cartilage degeneration of the LFC and LTP was divided into 4 types: type I was defined as all areas of the LFC or LTP were grade 0 of the ICRS classification; type II was defined as most of the degenerated cartilage area in the LFC or LTP was grade 1; type III was defined as most of the degenerated cartilage area in the LFC or LTP was grade 2; and type IV was defined as most of the degenerated cartilage area in the LFC or LTP was grade 3 or 4. Moreover, the medial compartment of the tibial plateau was subdivided into 3 zones using the ICRS cartilage injury evaluation.

### Table 1. The Abbreviations We Used in This Study.

| Measurement                          | Abbreviation | Subset      | Grade | Description                                                |
|--------------------------------------|--------------|-------------|-------|------------------------------------------------------------|
| Arthroscopy                          |              |             |       |                                                            |
| Anterior cruciate ligament           | ACL          | Intact      |       | Synovial sheath intact, no fiber disruption, solid insertion |
| Posterior cruciate ligament          | PCL          | Ruptured    |       | Synovial sheath not present, fibers attenuated             |
| Lateral meniscus                     | LM           | Intact      |       | Meniscus intact                                           |
| Medial meniscus                      | MM           | Degenerated |       | Meniscus degenerated                                      |
| Medial femoral condyle               | MFC          | Anterior    | Grade 0| Normal                                                    |
| Lateral femoral condyle              | LFC          | Central     | Grade 1| Superficial lesions                                      |
| Medial tibial plateau                | MTP          | Posterior   | Grade 2| Less than 50% of cartilage depth                          |
| Lateral tibial plateau               | LTP          |            | Grade 3| More than 50% of cartilage depth                          |
|                                      |              |            | Grade 4| Exposure of subchondral bone                              |
| Clinical                             |              | Varus       | Type A | Subchondral bone exposed central                           |
|                                      |              | Valgus      | Type B | Subchondral bone exposed anterior and central              |
|                                      |              |             | Type C | Subchondral bone exposed central and posterior             |
|                                      |              |             | Type D | Subchondral bone exposed from anterior to posterior        |
| Radiographic measurements            |              | FTA         | See Figure 2 |                                               |
| Femoral condylar femoral shaft angle | FCFS         | See Figure 2 |                                               |
| Femoral condylar tibial plateau angle| FCTP         | See Figure 2 |                                               |
| Tibial plateau tibial shaft angle     | TPTS         | See Figure 2 |                                               |

### Table 2. Assessments of the Anterior Cruciate Ligament (ACL), Posterior Cruciate Ligament (PCL), and Lateral Meniscus (LM) in Severe Varus Knee Osteoarthritis.

|                        | ACL          | PCL          | LM          |
|------------------------|--------------|--------------|-------------|
| Intact                 | 31 knees (44%) | 60 knees (85%) | 47 knees (66%) |
| Degenerated (LM) or ruptured (ACL, PCL) | 29 | 10 | 16 |
| Disappeared            | 11           | 1            | 8           |

*Intact: Visually as having an intact synovial sheath, no fiber disruption, and solid insertion points in the femur and tibia, and clinical assessment demonstrated a hard end point. Ruptured: The synovial sheath was not present, and fibers were visually attenuated or stretched, and when clinical assessment demonstrated a soft end point. Disappeared: The synovial sheath was not present and the fibers were visually absent.
package: anterior, central, and posterior. Based on the wear pattern of the MTP, severe varus knee osteoarthritis was divided into 4 types. The detailed information is shown in Figure 1.

The femorotibial angle (FTA) of the lower limb in the frontal plane was measured on standard preoperative weightbearing anteroposterior x-rays (Fig. 2). Four angles were then measured to assess the varus deformity of each knee in Figure 2.

The assessments of the ACL, PCL, and LM were compared with the radiographic alignment, and cartilage degeneration of the lateral compartment (LFC and LTP) was compared with the radiographic alignment. The type of MTP was also compared with the assessments of the ACL, PCL and MM, as well as the radiographic alignment. All data are expressed as mean and standard deviation, and statistical software (SPSS version 22, IBM, Armonk, NY, USA) was used for data analysis. Statistical analyses were performed using the Mann-Whitney U test (2 groups), Tukey-Kramer test (more than 2 groups), and the $\chi^2$ test. If the $P$ value was less than 0.05, the difference was considered significant.

**Results**

The assessments of the ACL, PCL, and LM in severe varus knee osteoarthritis are shown in Table 2. The APL normal group was defined as one in which the ACL, PCL, and LM were all intact in severe varus knee osteoarthritis, and the APL abnormal group was defined to include all other cases. With regard to each of the parameters of radiographic alignment, only the FTA of the APL normal group was significantly smaller than that of the APL abnormal group (Table 3). There were no significant differences in the other parameters of radiographic alignment. Therefore, the FTA of the patients who had normal ACL, PCL, and LM in varus knee osteoarthritis was significantly smaller than the others.

The comparison between the cartilage degeneration of the LFC and the radiographic alignment is shown in Table 4. The tibial plateau–tibial shaft (TPTS) angle was significantly
smaller in type I than in type IV. There were no significant differences in the parameters of radiographic alignment. Therefore, if the bone attrition in the medial tibia increased, the cartilage degeneration in the lateral femoral condyle increased.

The comparison between the cartilage degeneration of the LTP and the radiographic alignment is shown in Table 4.

The FTA was significantly smaller in type I than in type IV, and the TPTS was significantly larger in type IV than in types I and II. There were no significant differences in the other parameters of radiographic alignment in the cartilage degeneration of the LTP. Therefore, if the bone attrition in the medial tibia increased, the cartilage degeneration in the lateral tibial plateau increased.

Figure 2. Four angles to assess the varus deformity of each knee on standard preoperative weightbearing anteroposterior x-rays. (A) Femorotibial angle (FTA) between the anatomical axis of the femur and that of the tibia. (B) Femoral condylar–femoral shaft angle (FCFS) between the anatomical axis of the femur and the tangent to the subchondral plate of the femoral condyles. (C) Femoral condylar–tibial plateau angle (FCTP) between the tangents to the subchondral plates of the femoral and tibial condyles. (D) Tibial plateau–tibial shaft angle (TPTS) between the tangent to the subchondral plate of the tibia and the anatomical axis of the tibia.

Table 3. Comparison of the Femorotibial Angle Between the APL Normal and Abnormal Groups.*

| Mean (degrees) | Standard Deviation (degrees) |
|----------------|-----------------------------|
| 185.4          | 5.1                         |
| 188.4          | 5.6                         |

*The APL normal group was defined as the group in which anterior cruciate ligament (ACL), posterior cruciate ligament (PCL), and lateral meniscus (LM) were all intact in severe varus knee osteoarthritis, and the APL abnormal group was defined to include all other cases. There is a significant difference (P = 0.0400, Mann-Whitney U test). The figures in the parentheses indicate the number of cases.
The comparison between the assessment of the lateral meniscus and the radiographic alignment is shown in Table 5. The FTA was significantly smaller in type I than in type III, and the TPTS was significantly smaller in type I than in types II and III. There were no significant differences in the other parameters of radiographic alignment in the assessment of the LM. Therefore, if the bone attrition in the medial tibia increased, the degeneration of the lateral meniscus increased.

The details of the types of medial tibial plateau were shown in Table 6. There were no significant differences in the age at operation by medial tibial plateau type.
The ratio of intact ACL or PCL by the type of medial tibial plateau is shown in Figure 3. There was a significant positive correlation between intact ACL and type A medial tibial plateau ($\chi^2$ test; $P = 0.0096$). The ratio of surviving MM, including intact or degenerated, by the type of medial tibial plateau is shown in Figure 4. Independent of the type of medial tibial plateau, the middle and posterior segments of the MM disappeared more than the other portion of the MM. There were no significant differences in radiographic alignment by medial tibial plateau.

**Discussion**

In the present study, it appeared that, in the medial compartment, cartilage degeneration in the MTP first occurred in the central portion, and there was a significant positive correlation between intact ACL and type A MTP. Moschella et al.\(^5\) reported that the wear pattern in the MTP was limited mainly to the central portion in intact ACL knees and was observed to enlarge into the adjacent zone in ACL-deficient knees. Because the femur moves on the tibial plateau in a combination of rolling and sliding, and it has a wider contact area compared with the tibia; tibias presumably sustain constant mechanical stresses and, therefore, reflect the effect of mechanical stresses more than do femurs.\(^6\) Biomechanical descriptions of knee flexion and axial rotation during kneeling or squatting are consistent with the more pronounced anteromedial and posterolateral cartilage wear patterns observed in Saudi Arabian knees.\(^7\) The present results showed this in Japanese knees. There have been some reports of a correlation between subchondral bone and cartilage degeneration. It can be suggested that changes in histomorphometric parameters of subchondral bone are secondary to cartilage damage and proceed deeper into subchondral bone with increasing cartilage degeneration.\(^8\) Bone mineral density of the subchondral tibial bone is significantly correlated with future joint space narrowing and could be used as a predictor of knee osteoarthritis progression.\(^9\)

The ratio of intact ACL in patients with varus osteoarthritis undergoing total knee arthroplasty was reported to be 35.7%\(^5\) or 58.6%\(^10\); it was 44% in the present study. The ratio of disappeared ACL was reported to be 25%\(^7\) or 7.1%\(^10\); it was 15.5% in the present study. ACL rupture is not the primary etiology of the cartilage degenerative changes of the knee in patients affected by late arthritis.\(^7,11\) It seems that ACL rupture might be caused by late degenerative changes due to femoral intercondylar notch abrasion\(^12\) and by the increased joint shear force due to the absence of the physiological cartilage layer.\(^13\) Forty-six percent of PCLs from patients in the osteoarthritis group had moderate or marked degenerative change on light microscopy,\(^14\) but macroscopically, it was 15% in the present study.

The enlargement and displacement of the wear pattern in the MTP seems to be worsened by damage or absence of the MM.\(^15,16\) Moschella et al.\(^5\) found that severe cartilage wear in the central portion of the MTP was a median of 20% when the MM was intact, suggesting the occurrence of a protective function of the meniscus, even when the ACL was deficient. In the present study, the ratio of the middle segment of the MM was also the smallest in type A MTP. There was a strong correlation between the degree of MM subluxation and the severity of medial joint space narrowing.\(^17,20\) In the present study, independent of the type of MTP, the middle and posterior segments of the MM disappeared more than the other portion of the MM. This
Hunter et al. reported that cartilage loss was thought to be the major determinant of alignment, and that other factors, including meniscal degeneration and position, bone attrition, osteophytes, and ligament damage, contribute to the variance in malalignment. The knee bone mineral density distribution in the proximal tibia of the 113 patients was negatively correlated with the axial deformity. Increasing varus knee alignment was associated with increasing risk of medial compartment joint space narrowing and osteophytes. In the present study, if varus knee osteoarthritis progressed, the FTA, especially the TPTS, increased significantly. We thought that one of the main causes of progression of varus knee osteoarthritis was bone attrition in the MTP. Yamada et al. reported that the sclerotic changes associated with late-stage osteoarthritis were a late manifestation of the disease. Sclerotic changes probably reflected a response of the subchondral bone to the increased load bearing and friction that occurred after loss of the articular cartilage.

Varus thrust was a potent risk factor, identifiable by simple gait observation, for osteoarthritis progression in the medial tibiofemoral compartment. Ackroyd reported that the degenerative process usually started in the anterior half of the tibial plateau and in the corresponding part of the medial femoral condyle when the knee was in 20° to 30° of flexion. Buck et al. reported that osteoarthritis might not be a one-way road of cartilage loss. Subregional analysis suggested that, compared with healthy knees, cartilage changes in radiographic osteoarthritis might occur in both directions. Based on the results of the present study, the progression of varus knee osteoarthritis may occur in the following manner: Medial knee osteoarthritis starts in the central portion of the MTP, and, accompanied by medial meniscal extrusion and ACL rupture, cartilage degeneration expands from the anterior to the posterior in the MTP. Bone attrition occurs in the MTP, and the FTA and TPTS increase. Therefore, the lateral intercondylar eminence results in an injury of the cartilage of the longitudinal fissure type in the lateral femoral condyle, as shown in Figure 5. The cartilage degeneration then expands to involve the whole knee joints.

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Ethical Approval
This study was approved by our institutional review board.

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