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ABSTRACT: Mechanical factors have been implicated in the progression of knee osteoarthritis (OA). Understanding how these factors change as the condition progresses would elucidate their role and help in developing interventions that could delay the progression of OA. In this cross-sectional study, we identified kinematic and kinetic variables at the hip, knee, and ankle joints that change between three clinically distinct levels of knee OA disease severity: asymptomatic, moderate OA, and severe OA. The severity level was based on a combined radiographic/symptomatic clinical decision for treatment with (severe) or without (moderate) total knee replacement surgery. Gait variables that changed between groups were categorized as those that differed between the asymptomatic group and both OA groups, those that differed between the asymptomatic group and the severe OA group only, or those that changed progressively, that is, the asymptomatic differentiated from the moderate OA, and the moderate OA differed from the severe OA group. Changes seen in both OA subject groups compared to asymptomatic included increased midstance knee adduction moments, decreased peak knee flexion moments, decreased peak hip adduction moments, and decreased peak hip extension moments. Changes found only in the severe knee OA group included multiple kinematic and kinetic differences at the hip, knee, and ankle joints. Gait differences that progressed with OA severity included decreased stance phase knee flexion angles, decreased early stance knee extension moments, decreased peak stance phase hip internal rotation moments, and decreased peak ankle dorsiflexion moments. © 2007 Orthopaedic Research Society. Published by Wiley Periodicals, Inc. J Orthop Res 26:332-341, 2008

Keywords: knee osteoarthritis; disease severity; gait; biomechanics; lower extremity

INTRODUCTION

Knee osteoarthritis (OA) is a metabolically active, dynamic disease that includes both destruction and repair mechanisms that may be triggered by biochemical and mechanical insults. No definitive cure exists, and the pathomechanics are still not well understood, but in the past decade gait analysis has uncovered biomechanical characteristics of knee OA. Mechanical factors such as dynamic joint loading and mechanical loading have been implicated in knee OA pathomechanics, but their role in disease progression is equivocal. Most gait studies have characterized differences between subjects classified as either knee OA or asymptomatic. It is difficult to determine if the identified biomechanical changes were involved in the development of the disease, a response to degenerative changes in the joint and soft tissue, or a compensatory mechanism to the disease process. It is important to investigate how mechanical factors change as the condition progresses to understand the role of mechanical factors. While longitudinal studies could identify predictive factors, cross-sectional data from different levels of disease severity can provide information on which factors characterize each stage of the disease process.

Gait patterns of individuals with mild to moderate levels of knee OA have been investigated in an attempt to identify mechanical factors that are likely to be symptoms of pain and disability. In particular, the magnitude of the frontal plane knee adduction moment is higher than normal in those with moderate knee OA and relates to radiographic severity and short-term disease progression. However, a comprehensive description of the biomechanical changes associated with increasing levels of knee OA severity is lacking.
Knee OA presents a complex and multifaceted array of symptoms and risk factors, resulting in a very heterogeneous clinical condition that is difficult to classify. Most knee OA gait studies have classified subjects based on radiographic findings alone. However, there is little relationship between structural disease severity and symptoms. Clinically, a combination of radiographic and symptomatic factors is used to indicate disease severity and the need for total knee replacement, the common treatment for end-stage OA. In the present study, we used this clinical criterion to categorize disease severity, with subjects scheduled for knee replacement placed in a severe knee OA group. This assumes that candidates for knee replacement represent a distinctly different severity level than moderate knee OA patients, who were not identified as surgical candidates.

Our purpose was to identify kinematic and kinetic gait variables at the knee, hip, and ankle joints that distinguish between asymptomatic subjects and all subjects with knee OA (either moderate or severe), those that distinguish between asymptomatic and severe knee OA only, and those that change progressively from asymptomatic to moderate OA to severe OA. We hypothesized that many gait variable differences exist between the asymptomatic and severe OA groups, fewer between the asymptomatic and both OA groups, and fewer still that represent progressive changes between the three groups. From the results of previous gait studies, differences in the flexion angles at all joints, the flexion moment, the knee adduction moment, and hip adduction moment were expected.

**METHODS**

**Subjects**

The study included 60 asymptomatic subjects, 60 with moderate knee OA, and 61 with severe knee OA. Informed consent was obtained in accordance with the Dalhousie University Ethics Review Board. Asymptomatic subjects had no history of knee pain or surgical interventions to either lower limb and were recruited through a university news posting. They were screened for neuromuscular disorders, history of stroke and cardiovascular disease, and any other gait abnormalities.

Categorization into moderate and severe OA groups assumed that individuals indicated for total knee replacement represented a distinctly different clinical OA severity level than those who were not. Subjects indicated for joint replacement by a high-volume orthopedic surgeon within 6 months of the time of gait testing were categorized as severe; those who were not were categorized as moderate. The moderate subjects were recruited from the Orthopedic and Sports Medicine Clinic of Nova Scotia and selected from a waiting list for exploratory knee arthroscopy and from a list of individuals who had received exploratory arthroscopy surgery at least a year prior to the time of gait testing. These subjects were not candidates for knee replacement surgery. Subjects were excluded from this group if they had any major surgery or trauma to the lower limb, neuromuscular disorders, other forms of arthritis, gout, or history of stroke, or cardiovascular disease.

WOMAC self-reported pain, function, stiffness, and total scores were significantly greater in the severe group than the moderate and greater in the moderate group than the asymptomatic group (p < 0.0001). Kellgren–Lawrence (KL) radiographic scores, which assess joint severity by measurements of osteophyte presence, joint space narrowing, sclerosis, and joint deformity, were significantly different between the moderate and severe groups (p < 0.0001). The median KL score for the moderate group was 2, and scores ranged from 1 to 4. The median KL for the severe group was 3, and ranged from 3 to 4. All of the moderate subjects were either medial side dominant (medial side joint space narrowing less than lateral side joint space narrowing) or neutral (equal narrowing in both compartments). The majority (76%) of the severe subjects had predominantly medial side disease, but most showed degenerative changes in all compartments of the knee.

**Gait Analysis**

Three-dimensional (3D) motion of the lower limb and external ground reaction forces of all subjects were recorded with an Optotak 3020 optoelectronic motion capture system (Northern Digital, Inc., Waterloo, ON, Canada), and a synchronized AMTI force platform (Advanced Mechanical Technology, Inc., Watertown, MA). Three-marker triads of infrared light-emitting diodes were placed on the pelvis, thigh, shank, and foot body segments, and individual markers on the greater trochanter, the lateral epicondyle, the lateral malleolus, and the shoulder. Eight virtual markers were identified during quiet standing and used to define anatomical coordinate systems in each lower limb segment. Subjects were instructed to walk at a self-selected speed.

Intersegmental kinematics and kinetics were calculated by modeling the pelvis, thigh, shank, and foot as rigid bodies. The position and orientation of each segment at each time point in the gait cycle were computed using a least squares optimization routine. A complete gait cycle represents the time from first to second ground contact with the same foot. The 3D sign convention for the angles and moments at each joint followed a previously defined anatomically based coordinate system, where flexion/extension occurred about the medial/lateral axis of the proximal segment, internal/external rotation occurred about the distal/proximal or long axis of the distal segment, and adduction/abduction occurred about the floating axis (perpendicular to the flexion/extension and internal/external rotation axes).
Adduction/abduction and internal/external rotation angles were not examined because their relative values during gait are of a similar magnitude to that of the measurement error associated with kinematic cross-talk and skin motion.\textsuperscript{25-27} Net external intersegmental joint reaction moments were calculated using an inverse dynamics biomechanical model\textsuperscript{28} that combined positional data from the markers, ground reaction force and moment data, and limb inertial properties.\textsuperscript{29} All joint moments were normalized to body mass (kg).

**Statistical Methods**

The flexion/extension angles, the net external flexion/extension moments, abduction/adduction moments, and internal/external rotation moments at the hip, knee, and ankle joints were analyzed (12 gait measures in total). These measures were represented as waveforms that changed continuously throughout the gait cycle and were defined with 101 data points, one for each percentage of the cycle.

A total of 34 discrete parameters were extracted from the 12 gait measure waveforms. These parameters included peak, minimum, and range values chosen to characterize the shape of each waveform measure. For example, the knee flexion angle waveform characteristic shape included a stance phase peak flexion angle, a swing phase peak flexion angle, and a minimum value, so two peak values, a minimum, and a range value were analyzed. A one-way ANOVA was used to investigate differences in these parameters among the three subject groups. A Bonferroni correction for 34 comparisons at an \( \alpha \) level of 0.05 was used so that the effective \( p \) value for each analysis was set at 0.002 for significance. A Tukey post-hoc analysis was used for multiple comparisons among the groups. Results of the ANOVA and multiple comparison analyses were categorized into three groups for interpretation purposes: all OA gait changes, only severe OA gait changes, and progressive gait changes. All OA gait changes were those that differed between the asymptomatic group and both the moderate and severe groups. Gait changes that were only evident in the severe compared to the asymptomatic group or to the asymptomatic and moderate knee OA groups were considered only severe OA changes. Gait changes were considered progressive if they increased or decreased significantly between the asymptomatic, moderate knee OA, and severe knee OA groups.

**RESULTS**

Demographic, stride characteristic, and WOMAC differences among the three subject groups are summarized in Table 1. Similar to knee OA patient groups of other gait analyses, both the moderate and severe knee OA groups had higher body mass index (BMI) values than the asymptomatic group. The OA groups were older than the asymptomatic group. Speed decreased incrementally from the asymptomatic to the moderate knee OA to the severe knee OA group; stance percentage, stride time, and stance time all increased. Stride length was smaller in the severe OA group than the asymptomatic group. All WOMAC score subscales (pain, function, stiffness, and total) were higher in the moderate group than the asymptomatic group and higher in the severe group than the moderate group, indicating more severe self-reported pain, function, and stiffness.

Significant knee, hip, and ankle differences are summarized in Tables 2, 3, and 4, respectively. All OA differences at the knee joint included reduced early stance flexion moments (Fig. 1) and higher mid-stance adduction moments (Fig. 2). Both OA groups also had reduced peak and first peak hip adduction moments (Fig. 3), and reduced late stance hip extension moments. Only severe changes included reduced late stance knee extension moments (Fig. 1), reduced late stance knee internal rotation moments, and less range of sagittal plane joint motion at the knee, hip, and ankle joints. Progressive gait changes included two sagittal plane changes at the knee: successively smaller knee flexion angles during the stance phase (Fig. 4) and successively smaller knee extension moments in early stance phase (Fig. 1) from the asymptomatic group to the moderate OA group and from the moderate OA group to the severe OA group. Successively reduced late stance hip internal rotation moments and late stance ankle dorsiflexion moments were also progressive.

**DISCUSSION**

Several gait changes with knee OA have been reported.\textsuperscript{5,6,9-11} However, only a few investigators have attempted to capture how gait changes relate to disease progression, focusing primarily on knee adduction moment\textsuperscript{5,10,16,17} and, in a few cases, changes at the hip joint.\textsuperscript{12,13} In this study, we attempted to capture a more complete biomechanical description of lower extremity changes during gait that characterize different levels of disease severity. We hypothesized that the majority of differences would describe changes evident only in the severe knee OA group (only severe), but that some would characterize changes consistent with any level of severity (all OA), and some would display a trend that changes successively with severity level, suggesting the possibility of a link to a mechanism of disease progression (progressive).

As hypothesized, several only severe changes in gait variables were found, including reduced knee
Table 1. Subject Demographic, Stride Characteristic, and WOMAC Score Comparisons between the Three Subject Groups (A – Asymptomatic, M – Moderate OA, S – Severe OA)

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Group A Mean (±SD)</th>
<th>Group M Mean (±SD)</th>
<th>Group S Mean (±SD)</th>
<th>Multiple Comparisons</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td></td>
<td></td>
<td>M vs. A</td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td>25.48 (±4.04)</td>
<td>30.98 (±5.17)</td>
<td>32.05 (±5.48)</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Weight (kg)</td>
<td>73.5 (±14.19)</td>
<td>93.61 (±17.81)</td>
<td>91.08 (±15.92)</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Age</td>
<td>60.27 (±10.09)</td>
<td>58.32 (±9.31)</td>
<td>64.49 (±7.75)</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Speed (m/s²)</td>
<td>1.36 (±0.19)</td>
<td>1.25 (±0.22)</td>
<td>0.92 (±0.24)</td>
<td>0.002</td>
</tr>
<tr>
<td>Stride length (m)</td>
<td>1.44 (±0.13)</td>
<td>1.39 (±0.16)</td>
<td>1.16 (±0.19)</td>
<td>0.073</td>
</tr>
<tr>
<td>Stride time (s)</td>
<td>1.06 (±0.09)</td>
<td>1.19 (±0.12)</td>
<td>1.29 (±0.19)</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Stance time (s)</td>
<td>0.67 (±0.07)</td>
<td>0.73 (±0.09)</td>
<td>0.85 (±0.14)</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Stance %</td>
<td>62.8 (±1.59)</td>
<td>64.23 (±1.90)</td>
<td>65.64 (±2.26)</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>WOMAC pain</td>
<td>0.44 (±1.41)</td>
<td>7.53 (3.94)</td>
<td>10.62 (15.82)</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>WOMAC stiffness</td>
<td>0.31 (±1.02)</td>
<td>3.66 (1.66)</td>
<td>4.47 (1.76)</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>WOMAC function</td>
<td>1.67 (±5.01)</td>
<td>23.19 (±13.97)</td>
<td>34.37 (±17.8)</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>WOMAC total</td>
<td>2.27 (±6.9)</td>
<td>34.37 (±17.8)</td>
<td>50.89 (±17.7)</td>
<td>&lt;0.0001</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Gender</th>
<th>Group A Distribution</th>
<th>Group M Distribution</th>
<th>Group S Distribution</th>
</tr>
</thead>
<tbody>
<tr>
<td>Female</td>
<td>37 (62%)</td>
<td>40 (67%)</td>
<td>33 (54 %)</td>
</tr>
<tr>
<td>Male</td>
<td>23 (38%)</td>
<td>20 (33%)</td>
<td>28 (46 %)</td>
</tr>
</tbody>
</table>

There were significant group differences in all parameters using one-way ANOVA statistical analyses (p < 0.0001).

Multiple comparison p-values between subject groups for each parameter are shown.

Gender distribution among the subject groups is included at the bottom of the table.

Table 2. Knee Parameter Differences

<table>
<thead>
<tr>
<th>Gait Measure</th>
<th>Category</th>
<th>Group A Mean (±SD)</th>
<th>Group M Mean (±SD)</th>
<th>Group S Mean (±SD)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Knee abduction moment mid-stance minimum (nm/kg)</td>
<td>All OA</td>
<td>0.21 (±0.09)</td>
<td>0.29 (±0.13)</td>
<td>0.31 (±0.25)</td>
</tr>
<tr>
<td>Knee flexion moment peak (nm/kg)</td>
<td>All OA</td>
<td>0.52 (±0.29)</td>
<td>0.40 (±0.22)</td>
<td>0.33 (±0.20)</td>
</tr>
<tr>
<td>Knee flexion angle peak (°)</td>
<td>Only Severe</td>
<td>64.0 (±6.01)</td>
<td>61.3 (±7.86)</td>
<td>45.9 (±16.4)</td>
</tr>
<tr>
<td>Knee flexion angle range (°)</td>
<td>Only Severe</td>
<td>68.5 (±6.01)</td>
<td>66.0 (±7.40)</td>
<td>49.9 (±16.1)</td>
</tr>
<tr>
<td>Knee flexion moment late stance minimum (nm/kg)</td>
<td>Only Severe</td>
<td>-0.42 (±0.13)</td>
<td>-0.35 (±0.21)</td>
<td>-0.10 (±0.19)</td>
</tr>
<tr>
<td>Knee internal rotation moment peak (nm/kg)</td>
<td>Only Severe</td>
<td>0.21 (±0.053)</td>
<td>0.19 (±0.07)</td>
<td>0.12 (±0.08)</td>
</tr>
<tr>
<td>Knee flexion angle stance peak (°)</td>
<td>Progressive</td>
<td>18.72 (±7.28)</td>
<td>14.02 (±7.12)</td>
<td>8.04 (±6.22)</td>
</tr>
<tr>
<td>Knee flexion moment early stance minimum (nm/kg)</td>
<td>Progressive</td>
<td>-0.31 (±0.14)</td>
<td>-0.26 (±0.10)</td>
<td>-0.14 (±0.08)</td>
</tr>
</tbody>
</table>

Significant (p < 0.002) differences of one-way ANOVA on knee gait parameters are summarized, with Tukey post-hoc multiple comparison analyses.

Post-hoc analyses that indicated significant differences between both the moderate and severe OA subject groups with the asymptomatic group were considered All OA changes. Differences between the severe group and asymptomatic group (or both the asymptomatic and moderate groups) were categorized as Only Severe. Changes that increased or decreased incrementally from the asymptomatic to the moderate OA group and from the moderate OA group to the severe OA group were categorized as Progressive.
Table 3. Hip Parameter Differences

<table>
<thead>
<tr>
<th>Gait Measure</th>
<th>Category</th>
<th>Group A Mean (SD)</th>
<th>Group M Mean (SD)</th>
<th>Group S Mean (SD)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Hip adduction moment peak (mm/kg)</td>
<td>All OA</td>
<td>1.17 (±0.28)</td>
<td>0.99 (±0.26)</td>
<td>0.95 (±0.34)</td>
</tr>
<tr>
<td>Hip adduction moment early stance</td>
<td>All OA</td>
<td>1.15 (±0.28)</td>
<td>0.98 (±0.28)</td>
<td>0.95 (±0.34)</td>
</tr>
<tr>
<td>peak (mm/kg)</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Hip flexion moment late stance</td>
<td>All OA</td>
<td>−0.19 (±0.24)</td>
<td>−0.10 (±0.20)</td>
<td>−0.023 (±0.15)</td>
</tr>
<tr>
<td>minimum (mm/kg)</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Hip flexion angle range (°)</td>
<td>Only Severe</td>
<td>39.2 (±4.8)</td>
<td>39.8 (±5.1)</td>
<td>34.7 (±6.2)</td>
</tr>
<tr>
<td>Hip internal rotation moment</td>
<td>Progressive</td>
<td>0.11 (±0.09)</td>
<td>0.074 (±0.09)</td>
<td>0.031 (±0.04)</td>
</tr>
<tr>
<td>peak (mm/kg)</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Significant (p < 0.002) differences of one-way ANOVA on hip gait parameters are summarized, with Tukey post-hoc multiple comparison analyses.

Post-hoc analyses that indicated significant differences between both the moderate and severe OA subject groups with the asymptomatic group were considered All OA changes. Differences between the severe group and asymptomatic group (or both the asymptomatic and moderate groups) were categorized as Only Severe. Changes that increased or decreased incrementally from the asymptomatic to the moderate OA group and from the moderate OA group to the severe OA group were categorized as Progressive.

extension moments (Fig. 1) and reduced knee internal rotation moments in late stance phase, as well as smaller ranges of joint motion at all three lower extremity joints and reduced peak knee flexion angles (Fig. 4). Reduced knee extension moments in late stance have been associated with knee OA in previous studies. Messier and colleagues suggested that subjects with knee OA reduce the knee extension moment and therefore knee compressive forces by decreasing their walking speed in response to pain. This would substantiate finding this difference only in the severe knee OA population. Reduced range of motion is commonly reported as a response to the pain and dysfunction associated with degenerative joint disease, and reduced range of motion at all lower extremity joints has been reported. Individuals with more moderate disease generally have less pain and more mobility in their joints and therefore do not display the same deficiency in dynamic range of motion. Because the severe OA group consisted of individuals tested immediately prior to total knee replacement, these changes were more likely to be compensatory gait responses to pain and disability and therefore less likely involved in the pathomechanics of the disease.

All OA gait changes included reduced knee flexion moments in early stance (Fig. 1), which is consistent with previous studies, and higher mid-stance knee adduction moments (Fig. 2). Increased adduction moments are often associated with knee OA, but examination of peak adduction moments in previous studies has led to contradictory results. Our results support the recent suggestion that the mid-stance knee adduction moment is a more important parameter than the peak value for distinguishing between asymptomatic and OA gait patterns and the mid-stance knee adduction moment is a speed-independent measure, unlike the peak knee adduction moment.

An additional all OA change was reduced early stance peak hip adduction moments (Fig. 3).

Table 4. Ankle Parameter Differences

<table>
<thead>
<tr>
<th>Gait Measure</th>
<th>Category</th>
<th>Group A Mean (SD)</th>
<th>Group M Mean (SD)</th>
<th>Group S Mean (SD)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Ankle flexion angle range (°)</td>
<td>Only Severe</td>
<td>30.7 (±5.5)</td>
<td>30.3 (±5.0)</td>
<td>27.6 (±5.5)</td>
</tr>
<tr>
<td>Ankle flexion moment peak (mm/kg)</td>
<td>Only Severe</td>
<td>0.16 (±0.06)</td>
<td>0.15 (±0.06)</td>
<td>0.10 (±0.06)</td>
</tr>
<tr>
<td>Ankle flexion moment minimum (mm/kg)</td>
<td>Progressive</td>
<td>−1.39 (±0.11)</td>
<td>−1.26 (±0.18)</td>
<td>−1.1 (±0.18)</td>
</tr>
</tbody>
</table>

Significant (p < 0.002) differences of one-way ANOVA on ankle gait parameters are summarized, with Tukey post-hoc multiple comparison analyses.

Differences between the severe group and asymptomatic group (or both the asymptomatic and moderate groups) were categorized as Only Severe. Changes that increased or decreased incrementally from the asymptomatic to the moderate OA group and from the moderate OA group to the severe OA group were categorized as Progressive.
Figure 1. Knee flexion moment mean waveforms for three subject groups. Both OA groups had reduced stance phase knee flexion moments. Only the severe group had higher knee flexion moments in late stance as compared to the moderate and asymptomatic groups. Early stance phase knee extension moments were successively reduced from the asymptomatic to moderate group and from the moderate to severe group (all p < 0.002).

Figure 2. Knee adduction moment mean waveforms for three subject groups. Both OA groups had higher mid-stance knee adduction moments than the asymptomatic group (p < 0.002).
Mundermann and colleagues\textsuperscript{12} showed that more radiographically severe OA subjects had a lower peak hip adduction moment compared to less radiographically severe OA or control subjects. Chang and colleagues\textsuperscript{13} suggested that a greater peak external hip adduction moment was associated with a decreased likelihood of knee OA progression. Direct comparison with our results is difficult, as Mundermann and colleagues categorized groups by radiographs only, and the Chang study was an 18-month follow-up study. However, both of our knee OA subject groups had reduced peak hip adduction moments, consistent with the more radiographically severe knee OA group of Mundermann and colleagues, and consistent with a more radiographically progressed disease state in Chang’s study. Both OA groups also had higher late stance hip flexion moments, consistent with the results of Mundermann and colleagues.\textsuperscript{12}

Reduced stance phase knee flexion angles (Fig. 4) and early stance knee extension moments (Fig. 1) were identified as progressive. While previous studies reported smaller ranges of knee flexion angle during stance with knee OA,\textsuperscript{4,6} our study is the first to associate changes in both peak knee flexion angle and peak extension moment just after foot contact with increasing levels of knee OA severity. Previously, we had identified a multidimensional knee OA gait factor that represented a combination of gait differences during loading response that included the knee flexion moment.\textsuperscript{28} Further, while a few knee OA gait studies have investigated changes in other lower limb joints,\textsuperscript{7,12,13} none have reported the progressive differences in hip internal rotation moment or ankle flexion moment as shown in our data. These changes occurred in late stance and may reflect a response to the altered knee mechanics detected in early stance. Gait changes categorized as progressive may be important to the pathomechanics of knee OA and warrant further investigation.

Knee OA is a multifactorial disease process that involves many interrelated factors that interact to produce biomechanical changes throughout the disease process. The present study investigated lower limb gait changes associated with knee OA, but did not establish a hierarchy or explore relationships among the identified gait factors or with other important factors such as neuromuscular control, BMI, and walking velocity. For example, obesity is one of the most common risk factors of knee OA,\textsuperscript{26} and the OA subject groups had higher BMI levels that increased with severity (Table 1). In this study and others, joint moments were normalized to body mass to account for subject differences. As the majority of Canadians with knee OA are overweight or obese,\textsuperscript{27} it is difficult to control for BMI effects in the design of knee OA gait.
studies. Some previous studies have shown that obese individuals use altered gait mechanics and walk with increased total energy expenditure. However, some obese individuals may be able to reorganize neuromuscular function during gait, enabling them to maintain joint health, so the effect of obesity on joint mechanics and neuromuscular control warrants further study.

As in other knee OA studies, the subjects in the present study walked slower than the asymptomatic, and speed was reduced successively among the three severity groups (Table 1). Previous studies have demonstrated the effect of walking speed on gait measures, particularly in the sagittal plane. While gait speed causes biomechanical changes in asymptomatic individuals, no evidence exists of a clear causal path between gait speed and gait mechanics in the presence of knee OA. Gait speed is inherently linked to the disease process of knee OA, making it difficult to separate the two confounding effects. Some have attempted to account for the confounding effect of gait speed by selecting and comparing gait trials at a speed close to 1 m/s. However, this approach requires OA subjects to walk faster than they normally would, or asymptomatic subjects to walk slower than they prefer, or both. We wanted subjects to walk at their self-selected speed to characterize the natural mechanical environment of the joints during a daily activity. Another approach is to remove the effect of speed statistically with an analysis of covariance (ANCOVA) model. This was inappropriate in our study, because critical ANCOVA assumptions are violated when the covariate (speed) is affected by the treatment. ANCOVA is also contraindicated when large differences exist in the mean and range values of the covariate between groups, as shown in Table 1.

We extracted discrete parameters from the gait waveform profiles, but differences in the patterns of waveforms may provide more information on gait changes with knee OA severity. We previously developed methodology for extracting and comparing characteristic patterns of gait waveforms and for examining relationships between gait measures. Further study will use these techniques to identify multifactorial changes in gait waveforms that relate to disease severity. Finally, although we report differences among three clinically distinct subject groups, severity levels within each OA group covered a spectrum of disease both in terms of radiographs and symptoms. Investigation of how the biomechanical factors identified in this study relate to the spectrum of radiographic, pain, and functional measures should provide further insight to how these gait factors relate to disease progression.
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