



Abstract

 The relationship between age and gait characteristics in people with and without medial compartment osteoarthritis (OA) remains unclear. We aimed to characterize this relationship and to relate biomechanical and structural parameters in a subset of OA patients. Twenty five participants with diagnosed unilateral medial knee OA and 84 healthy participants, with no known knee pathology were recruited. 3D motion capture was used to analyse sagittal and coronal plane gait parameters while participants walked at a comfortable speed. Participants were categorized according to age (18-30, 31- 59 and 60+ years), and those with and without OA were compared between and within age groups. In a subset of OA patients, clinically available Computed Tomography images were used to assess joint structure. Differences in coronal plane kinematics at the hip and knee were noted in participants with OA particularly those who were older compared with our healthy controls, as well as increased knee moments. Knee adduction moment correlated with structural parameters in the subset of OA patients. Increased knee moments and altered kinematics were observed in older participants presenting with OA only, which seem to be related to morphological changes in the joint due to OA, as opposed to being related to the initial cause of medial knee OA. Key words: aging; gait; knee; osteoarthritis.

Introduction

 Knee joint osteoarthritis (OA) is one of the commonest diseases affecting the aging population, so there is a growing interest in the mechanisms underlying this degenerative disease. The progression of knee OA has been related to gait mechanics [\(1\)](#page-13-0), specifically relating to tempo-spatial parameters [\(2\)](#page-13-1). However, this can be complex since gait adaptations that occur in normal aging can be similar to those seen in pathological participants. A better understanding of normal and pathological aging would assist in the management of OA, if disease development or progression is directly related to certain characteristics of gait. The development of medial knee OA has been postulated to result from repetitive loading, leading to meniscal fatigue failure or acute injury, with subsequent medial compartment articular cartilage loss. This is supported by reports that OA incidence significantly increases with age [\(3,](#page-13-2) [4\)](#page-13-3), as well as evidence indicating that its incidence is associated with obesity, knee trauma and female gender [\(4\)](#page-13-3). In addition, an increase in external knee adduction moment (KAM), which is associated with increased medial knee- joint forces, has been reported in people with moderate to severe medial knee OA [\(5-7\)](#page-13-4), and has been correlated with disease progression [\(8\)](#page-13-5) and severity [\(9-11\)](#page-13-6). However, there is little evidence supporting the presence of higher KAMs in the early stages of knee OA [\(2,](#page-13-1) [11-13\)](#page-13-7), indicating higher medial loads may not be causative of medial knee OA. Recent studies have suggested that KAM can be influenced by

trunk motion [\(14\)](#page-13-8). People with knee OA demonstrate significant hip muscle weakness [\(15\)](#page-13-9), which has

been suggested to be a risk factor in the development of knee OA [\(16\)](#page-13-10) due to pelvic drop increasing

KAM magnitude [\(17\)](#page-13-11). Others however have proposed that hip abductor strength has little influence on

hip and knee adduction moments during gait [\(18,](#page-13-12) [19\)](#page-13-13).

 Changes in gait mechanics with healthy aging include decreased stride length and increased cadence [\(20,](#page-14-0) [21\)](#page-14-1), and associated alterations in sagittal plane hip and ankle kinematics and kinetics [\(20-22\)](#page-14-0). Altered knee flexion range of motion has also been proposed in healthy aging [\(22-24\)](#page-14-2), but changes in the coronal plane are less well understood. A slight increase in valgus static alignment has reported [\(25\)](#page-14-3) as well as increased abduction during early stance in gait [\(24\)](#page-14-4). Reduced medial knee-joint forces with advancing age have also been reported [\(22\)](#page-14-2). The association between aging, biomechanical loading, structural joint changes and OA development warrants further investigation, in particular coronal plane biomechanics due to their implication in knee OA. The aim of this study was to demonstrate the relationship between age and gait characteristics in people with and without medial compartment OA. Gait and structural parameters at the knee joint were also investigated in a subset of OA patients. We hypothesized that; i) younger OA subjects would have similar KAMs as healthy subjects; ii) older healthy subjects would have altered gait mechanics compared with young healthy subjects and; iii) older OA subjects would show deviations in gait parameters compared with healthy young and older subjects. Methodology This study had ethical approval from the South West London Research Ethics Committee and all participants provided written informed consent. Twenty-five participants with a clinical diagnosis of

unilateral knee OA affecting the medial compartment were recruited from Charing Cross Hospital and

local district regional hospitals, and 84 participants with no known knee pathology were recruited from

22 staff and students of Imperial College London. Participants were excluded from the study if they

demonstrated any neurological or musculoskeletal condition other than knee OA, rheumatoid or other

1 systemic inflammatory arthritis, morbid obesity (Body Mass Index  $>$  35 Kg/m<sup>2</sup>) or had undergone previous surgical treatment for knee OA. Gait Analysis Two Kistler portable force plates (Kistler Type 9286B, Kistler Instrumente AG, Winterthur, Switzerland) were embedded into a 6m walkway, and a 10 camera Vicon motion capture system (Vicon Motion Systems Ltd, Oxford, UK) were used to capture the position of reflective markers attached to the subject. Force plate data were recorded using an analogue signal data acquisition card provided with the Vicon system and the Vicon Nexus software at a sampling rate of 1000Hz. The motion capture was recorded at a rate of 100Hz and synchronised with the force data. Twenty reflective markers were positioned on the subject's pelvis and lower limbs and four marker clusters positioned on the subject's left and right thigh and calf segments [\(26\)](#page-14-5). Participants were asked to walk at a comfortable speed along the 6m walkway 5 times, or until three clean foot strikes had been recorded from each force plate. Kinematic and kinetic parameters were calculated using a custom model written in body builder software [\(26\)](#page-14-5). Joint moments were normalised to the subject's bodyweight x height. Data was separated into gait cycles for each leg based on vertical ground reaction forces and time-normalised. Hip, knee and 20 ankle joint angles at heel strike (HS), toe-off (TO) and the time point at which  $1<sup>st</sup>$  peak vertical ground reaction force (GRF) occurred were extracted using custom written code in Matlab. Only joint moments 22 at  $1^{st}$  peak GRF were extracted. The average of three trials was taken.

Sub-group Computed Tomography Analysis

 A subset of 6 of the OA participants additionally underwent preoperative CT scans of their affected knee using the Imperial knee protocol as part of their routine surgical preparation [\(27\)](#page-14-6). The scans were reconstructed (Robin's 3D Software) and femoral and tibial frames of reference were established. The radii of the medial and lateral flexion facets (LFF) were measured from spheres best fit to markers placed evenly across their surfaces, on the posterior aspect of the femur. To measure the radius of the medial extension facet (MEF), the femur was initially orientated so that its inferior aspect faced anteriorly and the flexion facets faced superiorly. The anterior and posterior borders of the MEF were then distinguished with markers located using the sagittal CT views. The posterior border defined the transition of the extension facet to the flexion facet, and the anterior border defined the transition from the extension facet to the trochlea. The bony surface between these two borders was subsequently also covered with markers and a sphere best fit to their positions. The radius of the sphere was determined as the MEF radius. The radii of the medial and lateral tibial plateaus (MTP and LTP, respectively) were measured 20mm below the most proximal aspect of the tibial spines. All measures were normalised to the medial flexion facet as this structure is relatively well preserved in OA.

## Statistical analysis

 Participants were separated into three age groups as follows; 18-30, 31-59 and 60+ years. Data from the left and right sides were averaged for healthy participants, and data for the affected and unaffected sides of OA participants were presented separately. All statistical analysis was carried out in SPSS (SPPS v21, IBM Corp, USA). Initially, multiple analysis of variance (MANOVA) was carried out to assess the 22 overall effects of age and OA group. Where the MANOVA attained significance multiple comparisons were carried out using one-way analysis of variance (ANOVA), two sample t-tests (OA versus healthy) and paired t-tests (OA unaffected versus affected sides). Linear regression was used to assess the

 relationship between structural measures and knee joint kinetics in the subset of OA participants that underwent CT imaging.

Results

 Healthy and OA subject details and spatio-temporal gait parameters for each age category are provided in Table 1. MANOVA revealed significant differences between OA and healthy participants for weight (p<0.01), walking speed (p<0.001) and stride length (p<0.05). There was no effect of age group on these parameters and no interaction (p>0.05). Post hoc analysis revealed that OA participants were significantly heavier, walked slower and with shorter stride lengths in the 60+ age category (Table 1). All kinematic and kinetic data have been presented in Tables 2 and 3. MANOVA for gait parameters revealed significant effects of both OA (p<0.001) and age (p<0.001) group, and significant interactions (p<0.005). Post hoc analysis revealed that coronal plane hip (p<0.001) and knee (p<0.05) angles and knee moment (p=0.001), and sagittal plane ankle moment (p<0.001) were affected by OA presence. Parameters effected by age group were coronal plane hip (p<0.05) and knee (p<0.01) angles and knee moment (p=0.002), and sagittal plane knee angle (p<0.05). Further statistical tests were carried out on these parameters only. .

*Coronal Plane Differences*

 All significant kinematic differences were found in the 60+ age group (Table 2). Hip adduction angle was significantly lower in OA participants affected side (at peak GRF) and significantly higher in their unaffected side (at TO), compared with healthy participants of similar age. At the knee, OA participants had significantly higher knee adduction angles at both HS and peak GRF compared with healthy



## *Spatio-temporal parameters*

 Older OA participants (60+ years) walked significantly slower and with a shorter stride length compared with healthy participants of a similar age; adaptations that are commonly reported among OA patients [\(2\)](#page-13-1), which may be a strategy to reduce pain and/or loading in the affected knee. Both kinematic and kinetic parameters are known to be affected by walking speed [\(28\)](#page-14-7), therefore the effect of altered speed should be acknowledged in interpretation of the results in the present study. There were no significant differences in walking speed in healthy older participants compared with the younger group, which conflicts with previous reports [\(29\)](#page-14-8). This may have been due to the relatively short walkway used in the present study (6m), possibly causing some healthy participants to walk at a lower velocity than their usual walking speed.

## *Coronal plane gait parameters*

 The main kinematic and kinetic impact of OA was noted in the coronal plane, with altered hip and knee adduction angles and higher KAM's in older OA participants. The magnitude of the increased KAMs are in agreement with several previous reports [\(5-7\)](#page-13-4), and have been correlated with both disease progression [\(8\)](#page-13-5) and severity [\(9-11\)](#page-13-6). Several mechanisms related to gait mechanics have been proposed to cause greater KAMs in OA participants. For example, a 'medial knee thrust', the abrupt appearance of varus alignment during the mid-stance phase of gait [\(30\)](#page-14-9), has been described previously in people with medial knee OA [\(30,](#page-14-9) [31\)](#page-14-10). However, this seems unlikely since the knee adduction angle appeared to be increased throughout the stance phase, as opposed to appearing abruptly in mid-stance.

 Alternatively, pelvic drop caused by contralateral hip abductor weakness has been proposed as a risk factor in medial OA progression [\(16\)](#page-13-10). We did note differences in coronal plane hip kinematics in our data, indicative of reduced hip ab/adduction range of motion in older OA participants. However, hip

 abductor weakness was unlikely to be the cause of higher KAMs in our data because OA participants actually showed greater hip abduction than healthy participants, and previous studies have reported no clear relationship between hip abductor muscle strength and specific KAM characteristics [\(18,](#page-13-12) [19\)](#page-13-13).

 As shown in Fig 1, these changes were only evident in the older OA participants; OA participants aged 31-59 years presented gait kinematics and kinetics that were similar compared with healthy participants of similar age and younger. Since the difference in walking speed between younger and older OA participants was not significant, it is unlikely that this result was due to differences in walking speed alone. It was notable that the increased KAM reflected an increase in lever arm rather than GRF in these participants, which is supported by the significant increase in knee adduction angle. The adaptations in coronal plane mechanics noted here, and in previous studies, do not appear to be causal of OA nor do they occur as a result increasing age in healthy participants; thus these adaptations likely occur as a result of increasing medial knee OA severity with increasing age. In support of this view, we previously reported no significant differences in KAMs between participants with early medial knee OA and healthy participants of a similar age during both gait [\(13\)](#page-13-14) and sit-to-stand tasks [\(12\)](#page-13-15). Increased KAMs in advanced OA may therefore relate to morphological changes in the joint, which have been reported to occur in knee OA including meniscus or cartilage wear [\(32\)](#page-14-11) with subsequent bone loss and other changes in the subchondral bone [\(33\)](#page-14-12). Loss of joint material ought to cause these individuals to generate higher KAMs. Altered cartilage volume and indeed subsequent erosion of bone in the wear scar of the weight-bearing regions of the knee joint, the consequential greater medial (relative to lateral) tibial bone area [\(34\)](#page-14-13) may result in altered (less balanced) load distribution between medial and lateral compartments.

 KAMs have also been associated with trunk motion [\(14\)](#page-13-8), and therefore alterations in the activation of the trunk musculature may contribute to increased KAMs in advanced OA. For example, weak hip musculature may cause pelvic drop [\(17\)](#page-13-11), increasing the KAM lever arm. Furthermore, an imbalance in medial-lateral activation of lower limb muscles (for example hamstrings or gastrocnemius) due to selective atrophy may impact medial knee displacement [\(35,](#page-14-14) [36\)](#page-14-15), affecting KAMs.

## *Sagittal plane gait parameters*

 The main sagittal plane changes related to increased age in our healthy participants. Older participants (60+) had significantly less knee extension compared with younger participants at HS, which is in agreement with previous work [\(22,](#page-14-2) [23\)](#page-14-16). Favre et al. [\(23\)](#page-14-16) proposed this was due to reduced tilt at the shank, attributed to reduced quadriceps strength with aging and a distal to proximal shift in power production [\(23\)](#page-14-16). Our data provide support for this proposal. Sagittal plane kinetic parameters were not affected by increased age in our healthy participants, which is in agreement with some previous work [\(23\)](#page-14-16). However others noted increased hip extensor moments and associated reductions in knee extensor [\(22,](#page-14-2) [37\)](#page-14-17) and ankle plantarflexor [\(22\)](#page-14-2) moments in elderly compared with young participants. These differences appear to become more pronounced with higher walking speeds [\(37\)](#page-14-17), and the study by DeVita & Hortobagyi [\(22\)](#page-14-2), which proposed a redistribution of joint torques during gait in elderly participants, asked participants to walk at a relatively fast speed (1.48m/s); a speed substantially greater than that adopted by older participants in the present study. Therefore, kinetic changes in healthy participants in the older age groups may be evident only at faster walker speeds. 

*Biomechanical vs structural parameters in OA participants*

23 In a subset of our knee OA patients, we noted significant correlations between KAM during gait and both 24 the medial tibial plateau and lateral flexion facet radii. Higher KAM at  $1<sup>st</sup>$  peak GRF was associated with a

 larger MTP radius. In agreement, a positive correlation between KAM impulse and medial tibial plateau bone area, and medial-lateral rotation of tibial bone size was previously noted in a larger group of OA participants [\(34\)](#page-14-13). These data support the theory that bone adapts structurally to cope with load and mechanical stress, and these changes may cause altered gait biomechanics. We also found a significant inverse correlation between KAM and LFF radius. It is possible that offloading of the lateral compartment also results in morphological adaptations on the lateral side of the joint; however, a larger study is necessary to corroborate this finding.

 There are two main limitations to this study, which should be acknowledged. Most importantly, we did not measure severity of OA. OA severity increases with age, however the age at which an individual develops OA may vary widely. In addition, healthy participants were recruited based on the criteria that they had no diagnosis of trauma or OA to the knee joints; therefore, we cannot rule out the possibility that some of these participants had early signs of knee OA, since they did not undergo imaging of the knee joint. However, we have previously shown that individuals with early OA have similar gait kinematics and kinetics compared with healthy participants [\(13\)](#page-13-14), therefore this possibility should not affect our interpretation of the data. It should also be acknowledged that the accuracy of kinematic and kinetic parameters in the lower extremities can be affected by soft tissue artefacts, resulting in high inter-subject variability particularly in the coronal plane [\(26\)](#page-14-5). This should be taken into consideration in the present data where only small kinematic or kinetic changes were found. However, a cluster-based model was utilised in this study, which has been shown to reduce the inter-subject variability of kinematics when compared with a more commonly used model (Duffell 2014 [\(26\)](#page-14-5).

 In conclusion, older participants with medial knee joint OA had significantly higher KAMs as well as altered coronal plane hip and knee kinematics. Increased KAMs were correlated with joint morphology.

- Altered gait parameters appear in older participants with OA only, which seem to be related to
- morphological changes in the joint due to OA.

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