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1 **Biomechanical and neuromuscular adaptations during the landing phase of a stepping-
2 down task in patients with early or established knee osteoarthritis.**

3

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34

35 **Abstract**

36 *Purpose.* To compare the knee joint kinematics, kinetics and EMG activity patterns during a
37 stepping-down task in patients with knee osteoarthritis (OA) with control subjects.

38 *Methods.* 33 women with knee OA (early OA, $n=14$; established OA $n=19$) and 14 female
39 control subjects performed a stepping-down task from a 20 cm step. Knee joint kinematics,
40 kinetics and EMG activity were recorded on the stepping-down leg during the loading phase.

41 *Results.* During the stepping-down task patients with established knee OA showed greater
42 normalized medial hamstrings activity ($p=0.034$) and greater vastus lateralis-medial hamstrings
43 co-contraction ($p=0.012$) than controls. Greater vastus medialis-medial hamstrings co-
44 contraction was found in patients with established OA compared to control subjects ($p=0.040$)
45 and to patients with early OA ($p=0.023$). Self-reported knee instability was reported in 7% and
46 32% of the patients with early and established OA, respectively.

47 *Conclusions.* The greater EMG co-activity found in established OA might suggest a less
48 efficient use of knee muscles or an attempt to compensate for greater knee laxity usually present
49 in patients with established OA. In the early stage of the disease, the biomechanical and
50 neuromuscular control of stepping-down is not altered compared to healthy controls.

51

52 **Key words:** biomechanics, stepping-down, osteoarthritis, muscle strength, knee instability.

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55

56 **1. Introduction**

57 Osteoarthritis (OA) is a highly prevalent joint disease [1], which has been counted globally as
58 the sixth leading cause of moderate-to-severe disability and the eight cause of burden disease
59 in the European region [2]. Patients with OA commonly experience pain, stiffness, reduction
60 in the range of motion and muscle weakness, factors associated with activity limitations such
61 as the difficulty to stand up from a chair, walk or climb stairs [3, 4]. Studies carried out in
62 patients with OA have documented the use of compensatory strategies during gait such as
63 decreased walking speed [5], decreased cadence [6], decreased stride length [7], decreased knee
64 flexion angle during the loading response phase[8], increased step width [9], increased hip
65 internal rotation and increased lateral trunk lean [9]. Modifications in knee loading distribution
66 such as increases in knee adduction moment (KAM) and knee adduction angular impulse have
67 also been reported [10, 11]. A direct association between higher KAM and severity of knee AO
68 has been found [10, 11]. Moreover, changes in EMG activity patterns during gait including
69 increased activity of hamstrings and increased co-contraction have been documented [12]. This
70 increased co-activation might be an adaptation of the individual with OA to deal with pain and
71 instability generated by the loss of joint integrity. In this view, this co-activation could increase
72 the stiffness of the joint promoting knee stability [9]. On the other hand, those gait modifications
73 and increased co-activation could interfere with the distribution of the load on the knee joint,
74 leading to further joint damage and disease progression [8].

75

76 The kinematic and kinetic characteristics during gait and stair climbing have been
77 extensively studied in patients with knee OA in comparison with healthy subjects [8, 13, 14].
78 However, analysis of the biomechanical characteristics involved in other activities of daily
79 living like stepping-down from a sidewalk still need to be further analyzed, as stepping-down
80 is a task that elicits complaints of instability and pain [15-17]. In addition some studies have

81 differentiated between the characteristics of patients in different stages of the disease (early vs.
82 established OA) but they often did not use MRI to define their groups. Knowledge of the stage
83 in the process in which modifications in movement patterns occur might be helpful in the
84 understanding of disease development and/or progression. It is possible that patients at risk or
85 with early OA, defined as joint pain with structural damage detected on MRI but hardly visible
86 on x-rays [18], respond better to certain interventions than patients with established OA.

87

88 Patients with knee osteoarthritis (OA) often complain of knee instability, defined as the
89 sensation of buckling, shifting or giving way, which usually translates into activity limitations
90 [19]. Previous studies have estimated that between 12% and 65% of this group of patients have
91 reported at least one episode of knee instability during the past three months [20, 21]. Knee
92 joint stabilization is thought to be influenced by active muscle force contraction and passive
93 ligaments restraints, both of which are usually affected in patients with knee OA [20, 22, 23].
94 Evidence has shown an association between self-reported knee instability and isokinetic
95 average knee muscle weakness [21], but not with passive knee laxity in this group of patients
96 [24]. However, failure to control the knee usually occurs during dynamic activities [19].
97 Therefore, in an attempt to further explore knee stability in patients with OA, recent studies
98 have aimed to identify the objective biomechanical and/or neuromuscular performance
99 characteristics associated with knee instability. Those studies have reported an association
100 between greater knee adduction moment and medial knee laxity during gait [10], and lower
101 medial knee muscle co-contraction prior to platform perturbations in patients with medial
102 compartment knee OA [25]. Nevertheless, to the best of our knowledge the biomechanical and
103 neuromuscular components associated with the sensation of knee instability in those patients
104 have not been fully recognized. In addition, assessment of self-reported knee instability has not
105 been specifically reported in patients with early OA. Further study of knee instability in patients

106 with early OA might help to clarify the association between knee instability and disease
107 severity. In knee OA, disease progression leads to a structural deterioration which subsequently
108 can cause joint instability, as often mentioned in OA. Nevertheless, joint instability can also
109 contribute to further disease progression [26].

110

111 During stair descent loading forces across the knee joints are higher than during stair
112 ascent and level walking, making it a more challenging task requiring good neuromuscular
113 control to obtain good shock absorption and knee stability [27, 28]. Particularly the early stance
114 phase is important during which the ground reaction forces need to be attenuated (by eccentric
115 muscle activity) as weight is loaded onto one limb [27]. Therefore, the stance phase of a step-
116 down task was assessed in the present study to represent the stance phase of stair descent. The
117 stepping-down task has been used successfully to study movement strategies in elderly subjects
118 [29] and dynamic knee instability in a patient with anterior cruciate ligament deficiency [16,
119 30]. Therefore, the purpose of this study was to investigate the joint kinematics, kinetics and
120 EMG activity patterns in patients with early or established OA of the knee during a stepping-
121 down task.

122

123 We hypothesize that the analysis of knee kinematics, kinetic and EMG activity during
124 the performance of the stepping-down task might elucidate relevant biomechanical
125 characteristics associated with compensatory strategies for instability or pain used by patients
126 with knee OA (early and established). Secondarily, this task might help to explore
127 biomechanical and neuromuscular strategies associated with self-reported knee instability in
128 this group of patients. The results might contribute to the design of intervention strategies
129 directed to treat difficulties of mobility and knee instability in patients with knee OA.

130

131

132 **2. Methods**

133 **2.1. Subjects**

134 A convenience sample of forty seven females was included in this study (Table 1). Patients with
135 OA ($n=33$) were recruited by a rheumatologist or orthopaedic surgeon from the University
136 Hospitals Leuven. Fourteen patients were classified as early OA based on a combination of
137 pain, Kellgren/Lawrance (KL) score=0 or 1 on radiography and presence of at least two of four
138 MRI criteria: (1) \geq BLOCKS grade 2 for size cartilage loss, (2) \geq BLOCKS for percentage full-
139 thickness cartilage loss, (3) signs of meniscal degeneration, and (4) \geq BLOCKS for size of
140 BMLs in any compartment [18]. Nineteen patients were classified as unilateral or bilateral
141 established knee OA based on the criteria from the American College of Rheumatology (ACR)
142 [31] and $KL\geq 2\pm$ [32, 33]. Control subjects ($n=14$) with no history of knee symptoms or
143 characteristics associated with knee OA and $KL=0$ were recruited from cultural or social
144 organizations. Demographic, clinical, radiographic, neuromuscular and biomechanical factors
145 related to OA were assessed. Total knee replacement, rheumatoid arthritis or any other form of
146 inflammatory arthritis (i.e. crystal arthropathy, septic arthritis, spondylarthropathy) were
147 considered exclusion criteria. All the participants provided written informed consent before
148 testing. The study was approved by the local Ethics Committee.

149

150 **2.2. Measures**

151 *2.2.1. Loading phase of stepping-down task.* The subjects were instructed to step down from a
152 wooden step (20cm) (Figure 1) onto a force plate with the evaluated limb and to step forward
153 with the other limb. Subjects ended in quiet stance on both legs in front of the force plate (Figure
154 2). The arms were kept flexed across the chest to avoid obstruction of the visibility of the
155 reflective markers. All patients wore standard sport shoes (kelme indoor copa). A task cycle
156 was considered from the first contact with the force plate (touch-down) until the toe-off from

157 the force plate with the evaluated limb. In a single session, three trials per patient were recorded.
158 Both limbs were assessed but only the index leg (see statistical analysis) was included in the
159 analysis.

160

161 *2.2.2. Knee instability.* Self-reported knee instability was evaluated based on a questionnaire
162 from Felson et al. [19, 20] in which a sensation of knee buckling, shifting or giving away during
163 the past 3 months was inquired. Persons reporting knee instability were additionally asked for
164 the number of episodes of instability, on which leg it was experienced. Knee instability was
165 dichotomized as “0” if they did not report episodes and “1” if they reported episodes of
166 instability during the past 3 months [18]. An additional question about history of knee injury
167 (“Did you ever have a knee injury?” yes/no) was formulated to persons who reported to have
168 had at least one episode of knee instability. This with the intention to explore whether the
169 sensation of instability could be due to another cause such as traumatic injury.

170

171 *2.2.3. Muscle strength.* Knee muscle strength was assessed using the Biodex System 3 Pro
172 (Biodex Medical System, Shirley, NY, USA). An initial practice attempt was used for the
173 participants to become familiar with the movements required. The patients performed 3
174 maximal test repetitions to measure the isokinetic strength of the knee extensor muscles (mainly
175 quadriceps) and knee flexor muscles (mainly hamstrings) for each knee, at 60°/s.” [34].
176 Isometric knee extension and flexion were measured in 60° flexion position. The peaks of three
177 trials were averaged in each leg separately for isometric and isokinetic assessments (quadriceps
178 and hamstrings torques (Nm)) and divided by patient’s weight (kg). This measure (in Nm/kg)
179 has excellent intra-rater reliability (ICC 0.93) in knee OA patients [35, 36].

180

181 2.2.4. *Knee joint alignment.* Knee alignment was measured from anterior-posterior weight
182 bearing radiograph of the lower limbs (Oldelft, Triathlon, Afga ADC M Compact Plus) by a
183 single experienced rheumatologist (FL). The alignment of the mechanical axis was reported as
184 varus if $\leq -3^\circ$ or valgus if $\geq 3^\circ$. Knee alignment between -3° and 3° was classified as neutral [37,
185 38].

186

187 2.2.5. *Activity Limitation.* Activity limitations were assessed subjectively using the Dutch
188 version of the Knee Injury and Osteoarthritis Outcome Score (KOOS) [39] which ranges from
189 0 (poor outcome) to 100 (good outcome), and objectively using the stairs test and the get up
190 and go test (GUG). In the stairs test [35], subjects were instructed to climb 5 stairs steps (15cm
191 high), turn around and descend the stairs. Participants were encouraged no to use the handrail,
192 but were not prohibited from doing so for safety. In the GUG test [35], subjects were sitting on
193 a high standard chair (49cm), they were told to stand up without help of the arms on the
194 command “go”, and walk 3 metres through an unobstructed corridor as fast as possible, without
195 running. Once they reached a mark on the floor, the subjects turned around, returned to the chair
196 and sat down. Patients who normally used walking devices were allowed to use them during
197 the test. All subjects were wearing standard sport shoes during the performance of the tests. The
198 time in seconds was recorded for both tests; longer time was considered a higher activity
199 limitation. For each test, the mean value of three trials was calculated. Both tests have shown
200 good reliability and validity [35].

201

202 2.2.6. *Pain and symptoms.* Pain was assessed with the visual analogue scale (VAS), the patient
203 was asked to range the sensation of pain during the last week from 0 (none) to 10 (severe pain).
204 The Dutch version of the KOOS questionnaire was also used to assess pain and general
205 symptoms, ranging them from 0 (poor outcome) to 100 (good outcome) [39].

206 2.3. Data capture
207 The stepping-down task was tracked using 6 MX-T20 optoelectronic cameras (Vicon, Oxford
208 Metrics, UK) collected at 100Hz in Nexus (Vicon). Eight body segments (trunk, pelvis, upper-
209 lower legs and feet) were identified by 46 spherical reflective markers of 14 mm diameter (see
210 Supplemental Digital Content from Malfait et al. [40] available at
211 <http://links.lww.com/MSS/A369>). Segmental coordinate systems were identified as reported
212 previously [41, 42]. Simultaneously (time synchronized), data from the force plate (AMTI
213 Watertown, MA, USA) and the electromyography (EMG) were sampled at 1500 Hz [34].

214

215 EMG activity of the vastus medialis (VM), vastus lateralis (VL), medial hamstrings
216 (MH) and lateral hamstrings (LH) was recorded bilaterally using a 16-channel system wireless
217 surface EMG system (Aurion, Italy) and silver-silver chloride, pre-gelled bipolar surface EMG
218 electrodes (Ambu Blue Sensor, Ballerup, Danmark). The electrodes were placed over the
219 muscle belly 2cm center-to center in line with the muscle fibres, and with an inter-electrode
220 distance of 3cm to reduce the possibility of cross-talk between neighbouring muscles [43].
221 Isolated manual muscle test [44] were used to validate the placement of the electrodes and to
222 assess for cross talk [45]. Skin surface was previously shaved and cleaned with 70% isopropyl
223 alcohol to reduce impedance.

224

225 2.4. Data processing and analysis

226 Separate trials were used for anatomical calibration and for calculation of hip and knee joint
227 centres and functional axis of the model [41, 46, 47]. Marker trajectories and force plate data
228 were both filtered using a 4th order low pass Butterworth filter with a cut off frequency of 20
229 Hz, based on previous studies [40, 48]. Touch-down and toe-off events were defined based on
230 the vertical force crossing a 20N threshold. Joint knee flexion angles were calculated at touch-

231 down and at the point of peak knee flexion during the task (PKFA) (Figure 3). Knee adduction
232 moment, defined as the external load applied at the joint moving the tibia to varus position was
233 calculated using inverse dynamics and normalized to body mass (Nm/kg). There were no clearly
234 defined early and late peak adduction moments during the performance of the stepping-down
235 task. Therefore, the peak knee adduction moment (PKAM) as well as the integral of the knee
236 adduction angular impulse (KAAI) over the complete stance phase (Nms/kg) were included in
237 the analyses. The average of 3 stepping-down trials was calculated for all biomechanical
238 parameters for each participant [49]. All modelling and analyses were undertaken in Visual 3D
239 (v.4.83, C-motion, Germantown, MD, USA) using geometric volumes to represent segments
240 based on cadaver segmental data as described in previous studies [40, 50]

241

242 EMG signals were high pass filtered at a cut-off frequency of 10 Hz [51]. The rectified
243 EMG signals were also filtered with a 4th order zero-lag low pass Butterworth filter at a cut-off
244 frequency of 50 Hz and subsequently normalized to the peak EMG activity of each muscle
245 during the stepping-down task cycle [52, 53]. The root mean square (RMS) from touch-down
246 to the PKFA was calculated for each muscle on the stepping-down leg.

247

248 Muscle co-contraction index (CCI) for the medial (VMMH=vastus medialis-medial
249 hamstrings) and lateral (VLLH=vastus lateralis-lateral hamstrings) sides of the knee joint, as
250 well as for the oblique surface of the knee joint (VLMH= vastus lateralis-medial hamstrings),
251 were calculated from touch-down to the PKFA according to the following equation [54]:

252

$$CCI = EMGS/EMGL \times (EMGS + EMGL)$$

253

255 in which EMGS is the normalized magnitude of the EMG signal for the less active
256 muscle and EMGL is the normalized magnitude of the EMG signal for the most active muscle.
257 To determine whether medial to lateral co-contraction was imbalanced muscle co-contraction
258 medial to lateral ratio was calculated dividing medial co-contraction index with lateral co-
259 contraction index [55].

260

261 The co-contraction index used here represents the balance of EMG activity between
262 pairs of antagonistic muscles and it is commonly used in the literature [55]. However, it is
263 important to consider that EMG signal does not reflect muscle force and hence this index does
264 not provide direct information about the magnitude of knee loading.

265

266 2.5. Statistical analysis

267 For the patients with knee OA an index knee was selected using the following decision tree: 1)
268 knee with established or early OA (ACR and KL score), if OA diagnosis was the same in both
269 knees, 2) instable knee and 3) painful knee. In participants in whom an index knee could not be
270 defined based on these signs, a random index joint was assigned. For the control subjects the
271 right knee was used as reference. The variables related to the index knee were used in the
272 analyses.

273

274 Descriptive statistics were used to characterise the study population, as well as the
275 patients with knee OA and control subjects separately. Percentages were used for categorical
276 variables, and means and SDs for continuous variables. ANOVA and chi-square tests were used
277 to analyse the differences in the distribution of the variables between the three subgroups.

278

279 One-way analyses of variance (ANOVA -Tukey post hoc tests) were used to test the
280 group difference in knee joint angles, external moments and EMG activity between subjects
281 with established OA, early OA and control subjects. Chi square tests (χ^2) were used to compare
282 self-reported knee instability between the study groups. Independent t-tests were used to
283 compare the patients' characteristics, joint kinematics, kinetics and EMG activity patterns
284 during a stepping-down task in patients with and without self-reported knee instability.
285 Statistical significance was accepted at p-values ≤ 0.05 . All analyses were performed using
286 SPSS software, version 17.0 (SPSS, Chicago, IL).

287

288 **3. RESULTS**

289 *3.1. Descriptives.* The mean age of the females that participated in the study was 68.9 (± 5.4)
290 years old. Patient with established knee OA had significantly more knee pain ($p=0.029$) and lower
291 isometric knee flexor muscle strength than the control group ($p=0.011$). A lower percentage of
292 patients with established OA had their knees in neutral alignment compared with patients with
293 early OA ($p=0.017$), the difference was borderline significant when comparing with control
294 subjects ($p=0.051$). No significant group differences were found in other variables assessed
295 including activity limitations. Further, demographic, clinical and neuromuscular characteristics
296 are shown in the Table 1.

297

298 *Knee biomechanics and EMG activity patterns during the loading phase of stepping-down task.*
299 There were no significant differences in kinematics or kinetics between the groups with knee
300 OA (early- established) and/or the control group during *the loading phase of the* stepping-down
301 task. Patients with established knee OA showed greater normalized medial hamstrings activity
302 ($p=0.034$) and greater vastus lateralis-medial hamstrings co-contraction ($p=0.012$) compared
303 with the control subjects. Higher vastus medialis-medial hamstrings co-contraction was found
304 in patients with established OA compared with control subjects ($p=0.040$) and to patients with
305 early OA ($p=0.023$) (Table 2) (Figure 4).

306

307 *3.2. Self-reported knee instability.* Seven patients (15%) with knee OA (early n=1; established
308 n=6) reported to have at least one episode of knee instability during the past three months. The
309 incidence of instability was significantly higher in the group with established OA compared
310 with the control ($p=0.020$). Neither of the patients with self-reported knee instability reported a
311 previous knee injury. None of the characteristics studied such as the biomechanics and EMG
312 activity patterns during the performance of the loading phase of stepping-down task (Table 3)

313 were significantly different between patients with or without self-reported knee instability.

314 However, patients with self-reported knee instability showed significantly lower knee muscle

315 strength compared with subjects without self-reported knee instability (Figure 5).

316

317 **4. DISCUSSION**

318 This study investigated the biomechanical and neuromuscular strategies during the loading
319 phase of a stepping-down task in a group of patients with early or established knee OA
320 compared to a healthy control group. The main study results showed no difference in the
321 kinematic or kinetic characteristics during the loading phase of a stepping-down task between
322 the three groups. However, greater muscle (co-)contraction patterns were observed in patients
323 with established knee OA compared with control subjects and patients with early OA.

324

325 There were no significant differences in sagittal plane kinematics or kinetics during the
326 loading phase of the stepping-down task between patients with early or established OA, and
327 control subjects. Based on these results, it is possible to conclude that an isolated stepping-
328 down task might not be challenging enough to identify kinematic and kinetic differences
329 between the three groups studied. Decreased knee flexion angle excursion was previously
330 reported in patients with established knee OA during a step down task from 20cm [8]. However,
331 in the present study, no significant difference in the knee flexion angle at touch-down, at peak
332 knee flexion during the stance phase or in flexion excursion was found between the three groups
333 studied during the loading phase of the stepping-down task. The difference between both studies
334 could be related to the fact that in the study carried out by Childs et al. [8] the subjects continued
335 to walk forward several steps after stepping-down, which may have allowed a more natural
336 performance of the task. The setting in our laboratory restricted the task only to one step forward
337 after stepping down (Figure 1).

338

339 Peak knee adduction moments and knee adduction angular impulse during the loading
340 phase of the stepping-down task were not significantly different between the three groups (early
341 OA, established OA and control subjects). Higher adduction moments during gait have been

342 previously found in patients with established knee OA in the medial compartment [10, 11, 56],
343 however not in subjects with early OA. It is therefore expected that knee OA severity in the
344 medial-compartment is associated with greater peak adduction moments. However, the
345 discrepancy with the results from the present study might be explained by the more
346 heterogeneous distribution of the structural features in the knee joint which is in line with
347 previous findings from Messier's et al. [57]. In addition, it is possible that the lack of association
348 between OA severity and knee adduction moment found during gait by other authors might not
349 be present during the loading phase of the stepping-down task evaluated in the present study.
350 Assessing the kinematics and kinetics of the supporting leg, in the step descent phase might
351 reveal more differences.

352

353 Greater medial hamstrings (MH) activity was exhibited in patients with established knee
354 OA compared with control subjects. Additionally, greater medial muscle co-contraction
355 (VMMH) was found in patient with established OA compared to control subjects and to patients
356 with early knee OA. These are in accordance with previous findings and may reflect an effort
357 to compensate higher medial knee laxity, usually present during gait in patients with established
358 OA [23-25]. Additionally, greater co-contraction of the posterior-medial (MH) and the lateral-
359 anterior (VL) side of the knee was found in the group of patients with established knee OA
360 compared with the control subjects. According to Rudolph et al. [54] high-level co-contraction
361 of opposing muscle groups could result in higher joint compression. These findings suggest not
362 only a higher medial compression of the medial knee compartment of the knee, but also an
363 overall increase in the compressive load through the knee surface in patients with established
364 OA. Previous evidence have suggested that an increase in muscle co-contraction may lead to
365 an increase of the cumulative load on the knee, which in turn might translate in further knee
366 joint damage and disease progression [8].

367 Seven patients with knee OA without a known history of knee injury reported to have
368 at least one episode of knee instability during the past three months. However, none of the
369 participants reported to have a feeling of knee instability during the performance of the
370 stepping-down task in our laboratory. In the present study, incidence of instability seems to
371 increase with the severity of the disease. However, to the best of our knowledge there is no
372 published evidence to prove this finding and the sample of patients with knee instability in this
373 study was too small to draw firm conclusion. It is possible that the study of biomechanical
374 characteristics of subjects with self-reported knee instability during the stepping-down task
375 might be useful to objectively identify performance characteristics associated with knee
376 instability, which could contribute to develop appropriate strategies oriented to counteract
377 instability in those patients. Nevertheless, probably due to the small number of patients with
378 self-reported knee instability within this study group, the results of this study did not support
379 our hypothesis. Therefore, studies in a larger sample population with self-reported knee
380 instability during the performance of a more challenging task might be needed to further clarify
381 whether or not biomechanical and neuromuscular performance based characteristics might be
382 associated with the feeling of instability in patients with knee OA.

383

384 In patients with established knee OA showing muscle weakness, muscle strength
385 training (both extensor and flexor knee muscles [34]) as well as neuromuscular training leading
386 to a selective EMG activity instead of increased and prolonged co-contraction patterns [58, 59]
387 may be recommended to preserve joint integrity (Hodges et al 2015). The influence of
388 neuromuscular training on knee stability still needs to be elucidated. Further studies are needed
389 to disentangle which of the biomechanical and neuromuscular performance based
390 characteristics are driven by pain, instability, structural changes and/or other factors. Overall,
391 it appears necessary to optimize the rehabilitation strategies directed to decrease an abnormal

392 joint loading during diverse activities of daily living in patients with OA. This might potentially
393 contribute to slow down the joint damage and subsequent increase in activity limitations in this
394 group of patients.

395

396 Several limitations of the present study should be considered. First, patients with uni-
397 and bilateral knee OA were included in the study. It is possible that patients with bilateral knee
398 OA might have developed different compensatory mechanisms to ambulate compared with
399 patients who have only one knee affected. However, it is very likely that all patients might have
400 had the contralateral knee (undiagnosed) affected to some extent. Overall, there is commonly
401 well-accepted to use an index knee, which includes the more affected knee in patients with
402 bilateral knee OA, for the analyses. Second, pain intensity during the performance of the
403 stepping-down task was not assessed. Authors are aware that pain could have influenced the
404 performance of the stepping-down task. Therefore, in a future study, gathering this information
405 will be considered in order to adjust the analyses. Third, only a small number of patients with
406 knee OA reported a sensation of knee instability during the past three months. The small number
407 of patients with this characteristic translated into a lack of statistical power which did not allow
408 us to draw strong conclusions about the biomechanical characteristics in patients with self-
409 reported knee instability from these analyses. Fourth, it was not possible to perform further
410 analyses by frequency of knee instability also due to the small number of patients with self-
411 reported knee instability. It is possible that patients with a higher number of episodes of knee
412 instability may have different biomechanical characteristics than patients with a lower number
413 of episodes. Therefore, self-reported knee instability should be used as an inclusion criterion on
414 for further studies in order to evaluate the kinetic and kinematic characteristics associated with
415 the sensation of knee instability. Fifth, differences in patients' height might have had a potential
416 influence on descending from a step [54]. However, there were no statistical differences in

417 height between the 3 study groups (Table 1). Additionally, a 20cm step is considered a standard
418 step height mimicking daily live scenarios involving stairs regardless of the height of the
419 patients. Lastly, assessing a flight of stairs rather than one step-down might have revealed more
420 differences.

421

422 **5. Conclusions**

423 The greater EMG activity found during the loading phase of the stepping-down task in
424 established OA might suggest a less efficient use of knee muscles or an attempt to increase knee
425 stability. Statistically significant differences in the other analyzed variables were not found

426

427

428 **Conflict of Interest**

429

430 The authors of this article declare that they have no conflict of interest.

431

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436

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FIGURES

Figure 1. Subject in the initial position

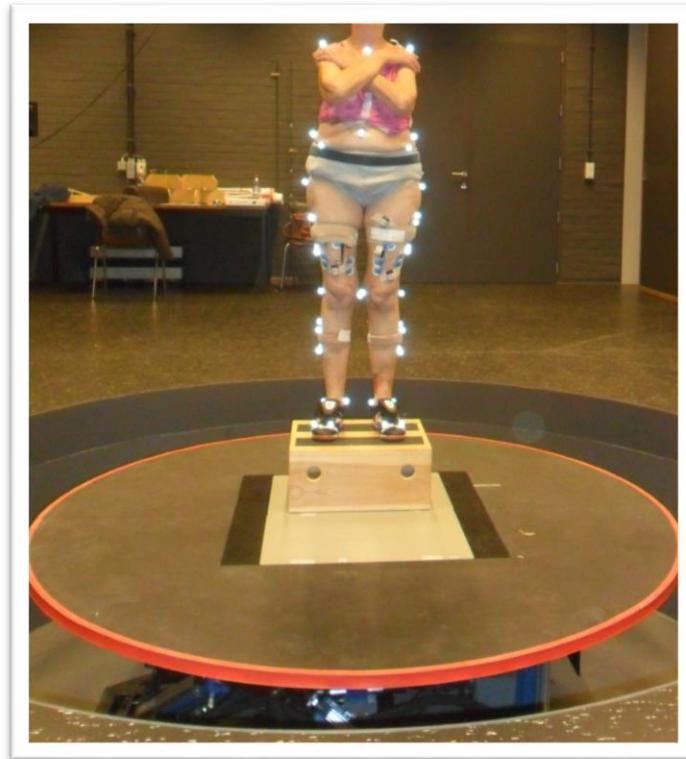


Figure 2. Stepping-down task

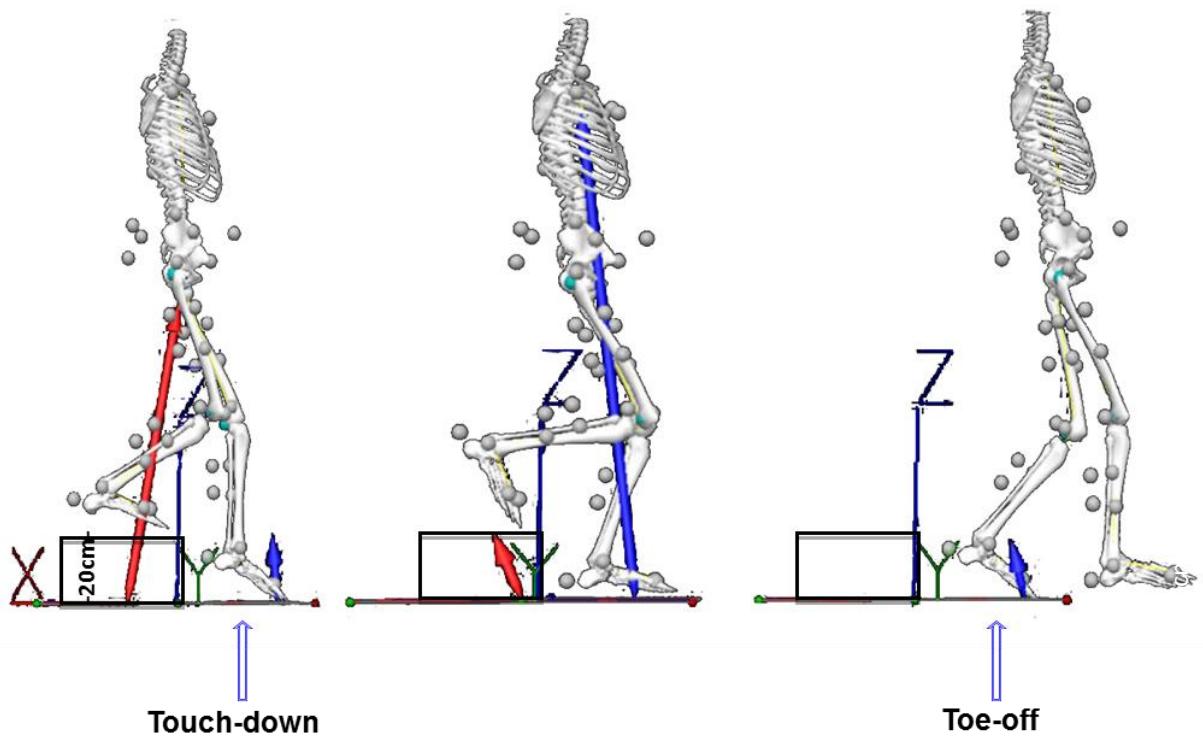
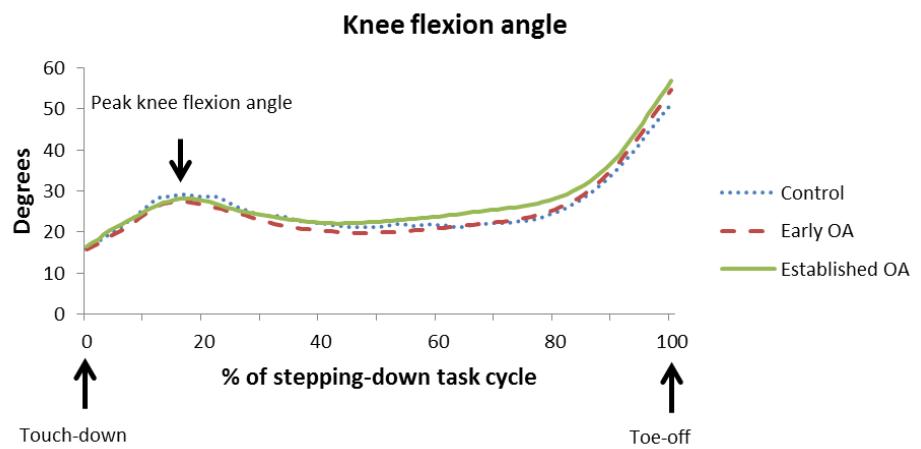


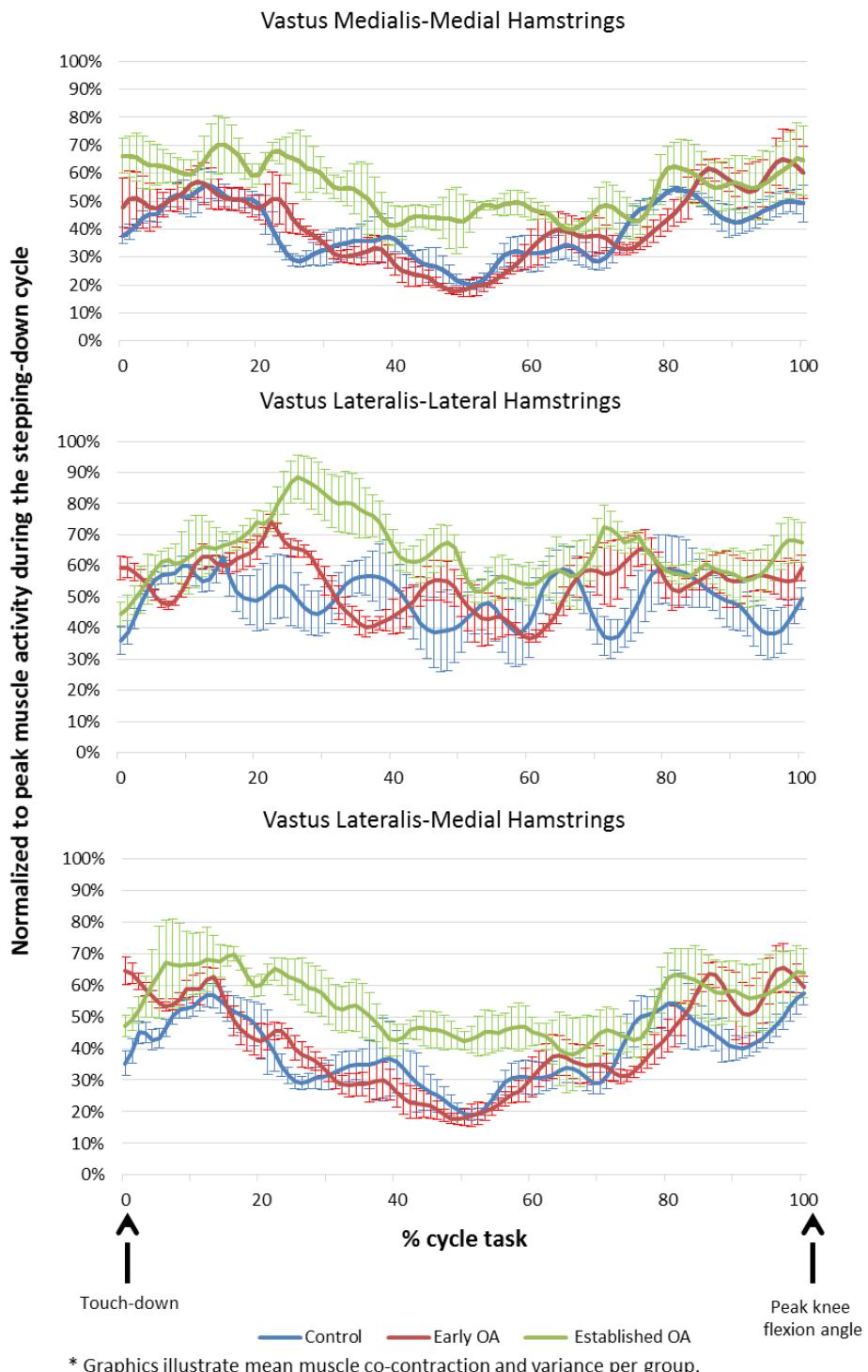
Figure 3



Lines represent the mean knee flexion angle per group.

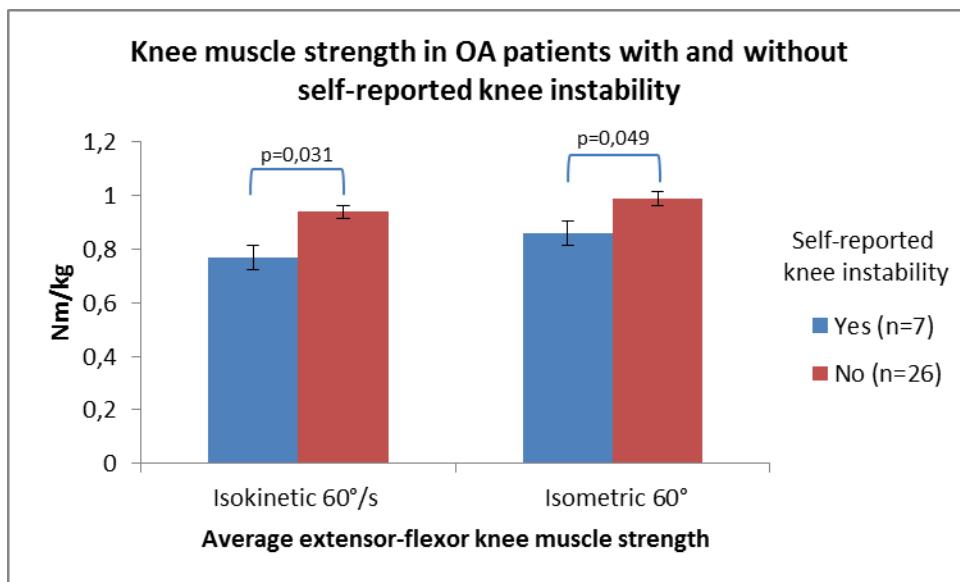
Figure 4

Muscle co-contraction during the stepping-down task



Muscle activity (EMG) of the loading leg was analyzed during the stepping-down cycle from the touch down (0%) to the peak knee flexion angle (100%). Lines represent the mean muscle co-contraction and variance per group.

Figure 5



TABLES

Table 1. Characteristics of study group

	Control subjects (n=14)	Early Knee OA (n=14)	Established Knee OA (n=19)	P- value	Post hoc P- value		
					Established vs control	Early vs control	Established vs early
Basic characteristics							
Age, in years	68.0±3.9	70.4±4.6	68.37±6.7	0.457			
Height, m	1.63±0.1	1.63±0.1	1.59±0.1	0.080			
Weight, kg	69.9±9.3	73.6±10.3	72.1±10.4	0.621			
Body Mass Index, kg/m ²	26.2±2.9	27.8±4.7	28.5±4.6	0.290			
K/L score, n (%)							
0	14(100)	-	-				
1	-	14(100)	-				
≥2±	-	-	19(100)				
Clinical characteristics							
VAS Knee Pain (0-10)	0.86±1.3	1.64±2.2	2.95±2.7	0.033*	0.029*	0.624	0.233
KOOS Pain score (0-100)	91.24±8.4	82.71±15.7	80.36±14.8	0.078			
KOOS Symptoms score (0-100)	89.50±10.1	78.77±15.4	74.96±18.2	0.033*	0.028*	0.163	0.764
Self-reported knee instability, n (%)	0(0)	1(7)	6(32)	0.026*	0.020*	0.309	0.090
Knee static alignment							
Varus (-) or valgus (+), degrees	1.06±2.1	-0.56±2.5	-0.61±3.6	0.216			
Neutral (>-3 and <3 degrees), n (%)	11(79)	12(86)	8(42)	0.027*	0.051	0.622	0.017*
Varus ≤ -3 degrees, n (%)	2(14)	1(7)	5(26)	0.164			
Valgus ≥3 degrees, n (%)	1(7)	1(7)	5(26)	0.337			
missing, n (%)	0(0)	0(0)	1(6)				
Muscle strength							
Isokinetic 60°/s							
Average knee muscle strength (Nm/kg) α	1.00±0.2	0.88±0.2	0.91±0.2	0.242			
Extensor muscle strength (Nm/kg)	1.20±0.3	1.01±0.2	1.08±0.3	0.227			
Flexor muscle strength (Nm/kg)	0.81±0.2	0.76±0.2	0.68±0.2	0.085			
Isometric 60°							
Average knee muscle strength (Nm/kg) α	1.10±0.2	0.99±0.2	0.95±0.1	0.078			
Extensor muscle strength (Nm/kg)	1.43±0.4	1.28±0.2	1.30±0.3	0.362			
Flexor muscle strength (Nm/kg)	0.76±0.1	0.68±0.2	0.62±0.1	0.015*	0.011*	0.217	0.445

Activity Limitations							
KOOS ADL score (0-100)	94.21±6.5	83.56±13.2	83.78±14.7	0.037*	0.058	0.066	0.999
Stair climbing test, seconds	5.67±1.1	5.50±1.1	5.94±1.0	0.510			
Get up and go test, seconds	6.53±1.7	6.28±1.4	7.09±1.6	0.338			
Cycle time stepping-down task §, seconds	1.07±0.2	1.09±0.2	1.04±0.2	0.734			

Mean ± standard deviation (sd), unless other stated. OA= osteoarthritis; K/L= Kellgren/Lawrence; VAS= visual analogue scale; KOOS= Knee Injury and Osteoarthritis Outcome Score; α Average knee extensor and flexor muscle strength; §Time from touch-down to toe-off. **Bold** χ^2 . * $p\leq 0.05$ significant difference between groups.

Table 2. Kinematics, kinetics and muscle activity during the stepping-down task

	Control (n=14)	Early OA (n=14)	Established OA (n=19)	P- value	Post hoc P- value		
					Established vs control	Early vs control	Established vs early
<i>Kinematics and kinetics</i>							
Knee flexion angle at touch-down, degrees	16.01±3.0	15.66±3.5	16.78±4.0	0.651			
Peak knee flexion angle (PKFA), degrees	31.01±6.1	29.02±4.2	30.59±6.7	0.636			
Knee flexion excursion, degrees	15.00±4.4	13.36±3.8	13.81±4.3	0.568			
Peak knee adduction moment (PKAM), Nm/Kg	0.37±0.4	0.29±0.1	0.30±0.3	0.697			
Knee adduction angular impulse moment (KAAI), Nms/Kg	0.23±0.3	0.16±0.1	0.17±0.2	0.598			
Peak knee flexion moment, Nm/Kg	-0.68±0.4	-0.65±0.1	-0.69±0.3	0.908			
Peak knee external rotation moment, Nm/Kg	-0.07±0.1	-0.07±0.1	-0.10±0.1	0.617			
<i>Muscle activity§</i>							
Vastus Medialis (VM)	0.42±0.1	0.41±0.1	0.43±0.1	0.720			
Vastus Lateralis (VL)	0.44±0.1	0.43±0.1	0.43±0.1	0.902			
Medialis Hamstrings (MH)	0.29±0.1	0.30±0.1	0.37±0.1	0.025*	0.034*	0.909	0.093
Lateral Hamstrings (LH)	0.33±0.1	0.36±0.1	0.39±0.1	0.298			
VMMH co-contraction	0.50±0.2	0.48±0.1	0.64±0.2	0.012*	0.040*	0.976	0.023*
VLLH co-contraction	0.55±0.2	0.64±0.2	0.65±0.2	0.310			
VLMH co-contraction	0.47±0.2	0.51±0.2	0.64±0.2	0.009*	0.012*	0.791	0.064
VMMH/VLLH co-contraction ratio	0.98±0.3	0.89±0.5	1.04±0.4	0.605			

§Root Mean square from touch-down to PKFA during the stepping-down task. Mean ± standard deviation (sd). * $p\leq 0.05$ significant difference between groups.

Table 3. Kinematics, kinetics and muscle activity during the stepping-down task in patients with knee OA (n=33) with or without Self-reported knee instability

Self-reported knee instability					
	Yes (n=7)	No (n=26)	p-value	r	p-value
<i>Kinematics and kinetics</i>					
Knee flexion angle at touch-down, degrees	15.12±5.4	16.62±3.2	0.355	-0.166	0.355
Peak knee flexion angle, degrees	30.09±8.2	29.88±5.1	0.951	0.015	0.934
Knee flexion excursion, degrees	14.97±3.5	13.26±4.2	0.331	0.174	0.331
Peak knee adduction moment (PKAM), Nm/Kg	0.28±0.4	0.30±0.2	0.874	-0.029	0.874
Knee adduction angular impulse moment (KAAI), Nms/Kg	0.22±0.1	0.15±0.2	0.249	0.206	0.249
Peak knee flexion moment, Nm/Kg	-0.73±0.4	-0.65±0.2	0.659	-0.125	0.490
Peak knee external rotation moment, Nm/Kg	-0.11±0.2	-0.08±0.1	0.705	-0.102	0.572
<i>Muscle activity</i> §					
Vastus Medialis (VM)	0.43±0.1	0.41±0.1	0.566	0.104	0.566
Vastus Lateralis (VL)	0.43±0.1	0.43±0.1	0.948	0.012	0.948
Medialis Hamstrings (MH)	0.36±0.1	0.34±0.1	0.507	0.120	0.507
Lateral Hamstrings (LH)	0.38±0.1	0.38±0.1	0.857	0.033	0.857
VMMH co-contraction	0.60±0.2	0.57±0.2	0.683	0.074	0.683
VLLH co-contraction	0.63±0.2	0.65±0.2	0.841	-0.036	0.841
VLMH co-contraction	0.60±0.2	0.58±0.2	0.834	0.038	0.834
VMMH/VLLH co-contraction ratio	1.04±0.5	0.96±0.4	0.684	0.073	0.684

§Root Mean square from touch-down to PKFA during the stepping-down task. Data are presented as mean ± standard deviation and r= Pearson correlation coefficient. No statistically significant differences between groups.

