BLOOD FLOW RESTRICTED WALKING IN ELDERLY INDIVIDUALS WITH KNEE OSTEOARTHRITIS: A FEASIBILITY STUDY

Naaja PETERSSON, MSc¹, Stian Langgård JØRGENSEN, MSc^{3,4,5}, Troels KJELDSEN, MSc^{2,5}, Inger MECHLENBURG, PhD^{2,5} and Per AAGAARD, PhD¹

From the ¹Department of Sports Science and Clinical Biomechanics, University of Southern Denmark, Odense, Denmark, ²Department of Orthopedic Surgery, Aarhus University Hospital, Aarhus N, Denmark, ³Department of Occupational and Physical Therapy, Horsens Regional Hospital, Horsens, Denmark, ⁴H-Hip, Horsens Regional Hospital, Horsens, Denmark and ⁵Department of Clinical Medicine, Aarhus University, Aarhus N, Denmark

Objective: To investigate whether blood flow restricted walking exercise is feasible in patients with knee osteoarthritis, and to examine changes in functional performance and self-reported function. **Design:** Feasibility study.

Patients and methods: Fourteen elderly individuals diagnosed with knee osteoarthritis participated in 8–10 weeks of outdoor walking (4 km/h, 20 min/ session, 4 times/week) with partial blood flow restriction applied to the affected leg. Adherence, dropouts and adverse events were registered. Timed Up and Go test, 30-s sit-to-stand performance, 40-m fast-paced walk speed, stair-climbing and Knee Osteoarthritis Outcome Score were assessed pre- and post-training.

Results: Nine participants completed the intervention, while 5 participants withdrew (4 due to intervention-related reasons). In non-completing participants baseline body mass index (BMI) (p = 0.05) and knee pain (p = 0.06) were higher, while gait performance (p = 0.04) was lower. Considering completed case data, the training-adherence rate was 93%, while mean knee pain in the affected leg was 0.7 on a numerical rating scale of 0–10. Functional performance improved, while self-reported function remained unchanged.

Conclusion: Blood flow restricted walking exercise appeared feasible in patients with knee osteoarthritis. Participants who completed the intervention protocol demonstrated improvements in functional performance, with no changes in selfreported function.

Key words: knee osteoarthritis; blood flow restriction exercise; physical functional performance; knee osteoarthritis outcome score; walking; ischaemia; occlusion; rehabilitation.

Accepted Feb 15, 2022; Epub ahead of print March 9, 2022

J Rehabil Med 2022; 54: jrm00282

DOI: 10.2340/jrm.v54.2163

Correspondence address: Naaja Petersson, Department of Sports Science and Clinical Biomechanics, University of Southern Denmark, Campusvej 55, DK-5230, Odense, Denmark. Email: Np_499@hotmail.com

LAY ABSTRACT

Patients with knee osteoarthritis typically experience knee joint pain and functional disability. Consequently, it is beneficial for these patients to engage in tolerable exercise programmes to enhance their functional performance. This study examined the feasibility of 8-10 weeks of blood flow restricted (BFR) walking exercise in patients with long-term knee osteoarthritis. The exercise programme was feasible, did not exacerbate knee pain or provoke cuff discomfort, and produced improved functional performance in those participants (9/14) who completed the intervention protocol (93% adherence rate). Notably however, 4 patients withdrew from the study due to intervention-related reasons. In this study, feasibility and adherence to BFR-walking appeared to depend on individual factors, with high body mass index, high perceived knee pain, and low baseline levels of fast-paced walking, predisposing to low adherence to training. Thus, these individual characteristics must be taken into account when administering BFR-walking exercise in patients with knee osteoarthritis.

K nee osteoarthritis (OA) is among the 10 most prevalent chronic conditions in Denmark (1). Patients with knee OA experience joint pain, stiffness and functional disability (2). Typically, knee extensor muscle atrophy and strength deficits (3, 4) occur in patients with knee OA, which has been associated with reduced functional performance (5, 6). Consequently, the loss of functional performance with knee OA may reduce patients' quality of life (QOL) (7–9). To increase or maintain functional performance and QOL, it becomes important to engage individuals with knee OA in exercise programmes that they can tolerate.

Heavy-load resistance training (HRT) (>70% of 1 repetition maximum) has been shown previously to improve lower limb muscle strength and functional performance in individuals with knee OA (10–12). However, the heavy exercise loads applied in HRT may be contraindicated in some individuals with knee OA due to exacerbated knee pain, which can limit adherence

to HRT intervention (10, 13). Thus, more tolerable, yet effective, exercise regimes are warranted for this particular population.

Low-load resistance exercise performed with concurrent blood flow restriction (BFR) typically achieved by means of pneumatic cuff compression applied proximally around the exercising limb has repeatedly demonstrated significant improvements in skeletal muscle mass, strength, and functional performance in both healthy, elderly individuals, and clinical populations (10, 14–19). BFR training is known to create a hypoxic myocellular environment in the working limb, which is suggested to give rise to elevated levels of metabolic stress that may lead to increased type II muscle fibre recruitment (20), myocellular swelling (21), and elevated intramuscular concentrations of metabolites (22) accompanied by increased synthesis of muscle protein. Furthermore, BFR-training has been observed to stimulate the proliferation of myogenic stem cells (satellite cells), which potentially improves the regenerative capacity of the muscle tissue and contributes to the marked muscle hypertrophy observed with this type of training (23). Concurrent cuff compression during walking (BFR-walking) has been shown to be a feasible exercise regime in healthy elderly individuals (19, 24). Furthermore, BFR exercise has been reported to cause substantial increases in lower limb muscle strength (6-22%) and muscle mass (3-5%) in response to 6-10 weeks intervention (14, 19). BFR exercise may also lead to marked increases in physical function in individuals over the age of 60 years (24).

Knee joint loading forces during horizontal walking are lower and thus potentially more tolerable to some OA patients than the loads applied with HRT. Consequently, the application of BFR-walking may represent a feasible and effective training modality for individuals with knee OA. Previous investigations have reported BFR exercise as safe when correctly and cautiously implemented (18, 25-29). However, only a single case study has investigated functional performance following BFR-walking in an individual with knee OA (30). Therefore, the aim of the current study was to investigate whether BFR-walking exercise is feasible in patients with knee OA in terms of adherence, drop-outs, adverse events, and knee pain responses to the training intervention. A secondary purpose was to examine changes in functional performance and self-reported knee joint function following BFR-walking exercise.

MATERIALS AND METHODS

Participants

Fourteen elderly individuals (70.4 ± 6.3 years of age) diagnosed with knee OA volunteered to participate in the feasibility study. The participants were recruited from a senior centre between 27 January 2020 and 30 January 2020. Individuals were considered eligible for participation if they were at least 60 years old, clinically diagnosed with knee OA, and radiographically verified. Exclusion criteria included: (i) previous knee replacement surgery (both knees), (ii) musculoskeletal or neurological diseases or impairments that could potentially be expected to hamper the execution of outdoor walking (4 km/h), (iii) previously experienced thrombosis events, (iv) cardiovascular problems without using prescribed medication for regulation of their conditions.

All participants were informed about the study procedures and signed a written informed consent in accordance with the Declaration of Helsinki. The Central Denmark Region Committee on Biomedical Research Ethics concluded that the study was exempt from ethical notification and approval (case number 1-10-72-148-19). The project was registered at the Danish Data Protection Agency (case number 1-16-02-51-20). Baseline characteristics of the study participants are listed in Table I.

Intervention

BFR-walking exercise was completed 4 times/week, with 1 supervised session and 3 non-supervised sessions each week. Training sessions consisted of 20-min outdoor walking performed at moderate walking speed (~4 km/h) with concurrent BFR applied to the affected leg (BFR-leg). In case of bilateral OA, the leg most severely affected by knee OA (based on the participant's experience with OA symptoms and disability) was chosen as the affected leg. Participants used either (i) the Endomondo App (Under Armour, Baltimore, Maryland) for smartphones to track time, speed and distance covered during exercise, or (ii) completed a predefined walking-route planned by the principal author (NP). Walking speed and session duration remained constant throughout the training period. In addition, the participants were instructed to continue their regular and habitual physical activities during the intervention period.

Before each training session, the participant placed the cuff around the most proximal portion of the thigh and inflated the cuff in a neutral standing position. During exercise the cuff pressure remained constant. BFR to the exercising limb was maintained during the entire 20-min walk. The pressure was released immediately after cessation of exercise. To ensure constant cuff pressure during walking, the participants were instructed to evaluate and adjust the occlusion pressure after 10 min of walking.

Due to restrictions during the COVID-19 pandemic, the supervised training sessions ceased early into the initiation of BFR-walking. A route diagram

	All participants $(n = 14)$		Intervention group $(n=9)$		Withdrawals $(n = 5)$		
Characteristics	Mean ± SD	[95% CI]	Mean ± SD	[95% CI]	Mean ± SD	[95% CI]	<i>p</i> -value
Sex, women, n	12		7		5		0.505
Age (yrs)	70.4±6.3	[67.1, 73.7]	70.4 ± 6.0	[66.5, 74.4]	70.4 ± 7.4	[63.9, 76.9]	0.990
Height (cm)	167±9	[162, 171]	170 ± 9	[164, 176]	161 ± 6	[156, 166]	0.075§
Body mass (kg)	85.8 ± 16.2	[77.3, 94.3]	83.7±18.1	[71.8, 95.5]	89.7±13.0	[78.3, 101.1]	0.527
BMI (kg/m ²)	31±5	[28, 34]	29 ± 5	[25, 32]	34 ± 4	[31, 38]	0.051 [§]
Disease duration (yrs)	6.1 ± 5.2	[3.4, 8.8]	5.9 ± 6.0	[1.9, 9.8]	6.5 ± 4.0	[3.0, 10.0]	0.836
Bilateral knee OA, n	10		6		4		1.000
AOP (mmHg)	240 ± 38	[220, 260]	229 ± 38	[204, 254]	260 ± 32	[232, 288]	0.149
60% AOP (mmHg)	151 ± 39	[131, 171]	138 ± 23	[123, 153]	176 ± 51	[131, 221]	0.075§
Physical function							
30STS (reps)	10 ± 2	[9, 11]	10 ± 2	[8, 11]	10 ± 3	[7, 13]	0.875
TUG (s)	9.7 ± 1.3	[9.0, 10.4]	9.3 ± 1.4	[8.4, 10.3]	10.2 ± 1.1	[9.3, 11.2]	0.237
40MWT (m/s)	1.48 ± 0.18	[1.39, 1.58]	1.55 ± 0.13	[1.47, 1.64]	1.36 ± 0.19	[1.19, 1.53]	0.044*
11-step SCT (s)	16.3 ± 6.5	[12.9, 19.7]	13.9 ± 2.4	[12.3, 15.4]	20.8 ± 9.4	[12.5, 29.0]	0.181
KOOS (score 0-100)							
Pain	65 ± 19	[56, 75]	72 ± 15	[62, 82]	53 ± 19	[37, 69]	0.057⁵
Symptoms	77±17	[68, 86]	80±19	[67, 92]	73 ± 14	[61, 85]	0.500
ADL	67±20	[56, 77]	76±12	[68, 84]	50 ± 22	[31, 69]	0.053§
Sport	28 ± 24	[15, 40]	34 ± 26	[43, 56]	15 ± 13	[4, 26]	0.149
QOL	44 ± 17	[35, 52]	49 ± 10	[43, 56]	33±23	[13, 53]	0.179

BMI: body mass index; Yrs: years since diagnosis; AOP: Arterial occlusion pressure; 30STS: 30 seconds chair-stand test; TUG: Timed Up and Go; 40MWT: 40m fast-paced walk test; 11-step SCT: 11-step stair-climb test; KOOS: Knee Injury and Osteoarthritis Outcome Score; ADL: activities of daily living; QOL: quality of life. p-values represent comparisons between the Intervention group and the Withdrawal group. *p < 0.05, *p < 0.10.

depicting the individual intervention activities is shown in Fig. 1.

Blood flow occlusion

Arterial occlusion pressure (AOP) of the affected leg was determined with participants seated on an examination table in a resting state. An 11-cm wide pneumatic cuff



Fig. 1. Overview of patient flow. Due to rolling recruitment and baseline testing of participants, each cluster of participants (Groups 1–3) began blood flow restricted (BFR)-walking exercise in different weeks.

(Occlude Aps, Aarhus, Denmark) was placed around the proximal portion of the participant's thigh on the BFR-leg. To determine AOP an Ultrasound Doppler transducer (Edan SD3 Vascular Ultrasonic Pocket Doppler, Edan, USA) was placed posterior to the medial malleolus on the affected lower limb to identify the arterial pulse in the tibial artery. When the pulse was identified, the cuff was incrementally inflated up to the point where the peripheral arterial pulse was eliminated. The pulse elimination pressure would be determined as the AOP. For safety reasons, in case the pulse was still present at 300 mmHg, the AOP was set to 300 mmHg. Subsequently, an individualized cuff pressure of 60% of AOP was applied for each participant during BFR-walking (24, 31–33).

Outcome measures

All participants underwent baseline testing 1 week prior to the first exercise session. Participants were retested (post-testing) after 8–10 weeks of BFR-walking exercise (4 times weekly), in the week following completion of their final training session. Test procedures are elaborated below.

Feasibility and adherence

After each training session, participants registered total walking-time into a designated training diary, along with distance covered, cuff pressure, knee pain in the BFR-leg during walking (numerical rating scale (NRS) 0–10, scores of 10 representing maximal intolerable pain), and perceived exertion in the BFR-leg during walking (NRS 0–10, score of 10 representing maximal and complete exertion). Also, the participants were instructed to report any adverse events or complica-



Fig. 2. Setup of (A) 30 seconds chairstand test (30STS), (B) Timed Up and Go (TUG) test, (C) 40m fast-paced walk test (40MWT), and (D) 11-step stair-climb test (11-step SCT).

tions related to the cuff application during training. Adherence was calculated as:

Total training sessions completed (number of sessions)	
Total weeks completed (number of weeks)	*100
Veekly training sessions scheduled to perform (number of ses	sions)

= adherence (%)

Functional performance

Physical performance was assessed before and after the intervention period using the 30-s chair sit-to-stand test (30STS), Timed Up and Go (TUG), 40-m fastpaced walking (40MWT) and stair-climbing (11-step SCT) (Fig. 2). All tests are characterized as reliable and valid performance tests (5, 34). A description of each test including equipment and procedures are described by Tolk et al. and Dobson et al. (34, 35). The tests were intended to assess functional and balance performance and lower body strength, and have all been extensively used to test individuals with knee OA (5, 6, 24).

Self-reported knee function

Knee joint pain, knee-related symptoms, function in daily living (ADL), function in sport and recreation and QOL were assessed using the Knee Injury and Osteoarthritis Outcome Score survey (KOOS). The KOOS questionnaire has been demonstrated as a reliable and valid tool for evaluating perceived knee function and associated problems in persons with knee OA (36, 37).

Statistical analysis

Microsoft Excel (version 16.36, Microsoft Corporation, USA) and SPSS statistics (version 26, IBM Corporation, USA) were used to compute the data. The Shapiro–Wilk test was performed to assess if data were normally distributed. Descriptive statistics are presented as group means±SD and 95% confidence interval (95% CI) unless otherwise stated.

To examine any changes in baseline data between the group of participants who completed the intervention period (n=9) and the withdrawals (n=5) an independent *t*-test or the Mann–Whitney *U* test was applied, depending on whether the data was normally distributed. Furthermore, changes from pre- to post-intervention, as well as development in knee joint pain, and perceived exertion during BFR-walking exercise, were examined using paired *t*-testing. Wilcoxon signed-ranked testing was used in cases where data did not follow a normal distribution.

RESULTS

Baseline characteristics

Nine participants completed the BFR-walking training protocol and were evaluated at post-testing, while 5 participants withdrew from the study during the intervention period. A number of differences were noted between the 2 groups (Table I). Baseline measurements demonstrated that participants who withdrew from the study showed: *(i)* slower walking speed during the 40MWT (p=0.044), *(ii)* a tendency to elevated perceived knee pain (p=0.057), *(iii)* a tendency to higher body mass index (BMI) (p=0.051), and *(iv)* a tendency to worse self-reported ADL (p=0.053) in comparison with participants who completed the intervention protocol.

Feasibility and adherence

In total, 8–10 weeks of BFR-walking exercise was performed by 9 participants. Six participants completed 9 weeks training, 2 participants completed 10 weeks of training, and only 1 participant completed 8 weeks of BFR-walking training. They completed a mean of 34 ± 4 training sessions per participant, and the mean±SD adherence to the training sessions was 92.8 ± 9.2 % considering completed case data.

The mean \pm SD (min-max range) cuff pressure during BFR-walking was 137±23 (96–173) mmHg. Target cuff-pressures were maintained throughout the study period in 7 of the 9 participants, but had to be reduced in 2 participants (to reach levels corresponding to $\sim 50\%$ AOP) due to lower limb discomfort during BFR-walking. The first participant experienced discomfort in the lower leg due to varicose veins, which, according to a consultant vascular surgeon assessment, was not deemed dangerous or harmful to the person. However, discomfort in the lower leg did occur, and thus the cuff pressure was reduced for the final 2 weeks of exercise. The second participant experienced a leg discomfort of unknown cause in the last 2 weeks of exercise, which could not be related directly to the BFR-walking training. Despite 2 participants experiencing discomfort from the cuff perssure, the mean \pm SD perceived exertion in the BFR-leg was rated 3.4 ± 2.1 on the 0–10 point NRS.

The mean \pm SD knee joint pain during BFR-walking was 0.7 ± 0.9 on the 0–10 point NRS.

Drop-outs

Five participants withdrew from the study during the intervention period due to; (*i*) exercise-induced discomfort when using the cuff (n=3), (*ii*) exacerbated knee pain during the BFR-intervention period (n=1), (*iii*) pain in the contra-lateral knee joint (n=1) (unknown relation to BFR-walking exercise). It should be noted that 1 of the participants who withdrew from the study due to limb discomfort was excluded from the study, since she could not manage to safely apply the cuff at home according to the desired method in the current study.

Functional performance

Signs of improved functional performance were observed after the period of BFR training (Fig. 3) (p < 0.05) for 30STS (+16%), TUG (-8%) and 40MWT (+5%) (Table II). In contrast, the 11-step SCT test remained unchanged (+0.7% n.s.) (Table II).

Self-reported knee function

KOOS subscale scores remained unchanged following the period of BFR-walking (Table II).

DISCUSSION

This is the first study to examine the feasibility of BFR-walking exercise in elderly individuals with knee OA. The main findings were that participants who completed 8–10 weeks of BFR-walking exercise demonstrated: (*i*) high adherence to training, (*ii*) minimal knee pain and exertion during BFR-exercise, (*iii*) significant improvements in functional performance (30STS, TUG and 40MWT); however, (*iv*) without demonstrating any changes in KOOS subscale scores. Notably, 5 participants withdrew from the study, of which 4 participants experienced intervention-related adverse events, such as exacerbated knee pain in the BFR-leg or discomfort from the cuff during BFR-walking.

Feasibility and adherence

Participants who completed the intervention period demonstrated an exercise adherence of 93% to the scheduled training sessions, which is in line with previous study reports in comparable populations (OA) and exercise conditions (resistance training intervention with and without BFR) (10). Furthermore, perceived knee pain-scores and exertion in the BFR-leg were low (NRS 0.7 and 3.4, respectively). These results indicate that the application of a pneumatic cuff during the exercise was tolerable for the majority of the present group of OA patients and did not contribute to exacerbated knee pain nor lead to increased levels of discomfort.

However, 4 participants experienced adverse events during the BFR-walking intervention period, which led to withdrawal from the current study. A number of differences existed between the participants who completed the intervention period and participants withdrawing from the study (cf. Table I). These differences might have been, at least in part, contributing to the observed withdrawal of participants. Thus, it could be speculated that the extent of knee pain may have negatively affected the adherence to exercise. Furthermore, differences such as slower 40-m walking speed and a tendency to lower self-reported ADL were found in withdrawing compared with completing participants, which may have affected at least some participants' ability to complete the intervention period. Thus, walking continuously for 20 min may be difficult for some patients with knee OA. Alternatively, BFRwalking, performed as either interval-based training or simply reducing the walking time to 10 min have previously been demonstrated to improve functional performance (24). These BFR-walking exercise protocols would perhaps increase the overall adherence to the study.

As reported by Gomes-Neto et al. (2015) obese elderly people with knee OA demonstrate reduced functional



Fig. 3. Functional performance assessed before and after 8–10 weeks of blood flow restricted (BFR)-walking exercise in elderly patients with knee osteoarthritis (OA). Pre and post measurements in the 30-second chair sit-to-stand test (30STS), Timed Up and Go (TUG) test, 40-m fast-paced walking (40MWT), and 11-step stair-climbing test (11-step SCT). Error bars denote SD. *p < 0.05, Pre vs Post.

performance, higher levels of knee pain and greater difficulty in performing everyday tasks compared with non-obese age-matched elderly people with knee OA (38). Thus, it is possible that a higher BMI, as seen in the current group of withdrawing participants, might have complicated their ability to complete the BFR-walking exercise.

Notably however, a majority of the current participants experienced BFR-walking as a tolerable and feasible exercise paradigm, which could be performed with a relatively high exercise adherence and with minimal knee pain and discomfort. Nonetheless, the current data suggest that individuals with high BMI and high degrees of perceived knee pain and low ADL function may benefit less from this type of exercise, as manifested by events of exacerbated knee pain and discomfort from the occlusion cuff.

 Table II.
 Functional performance and Knee Injury and Osteoarthritis Outcome Score (KOOS) subscale scores pre-and-post 8–10 weeks

 of blood flow restricted (BFR)-walking exercise

	Pre-test		Post-test		Change		
	Mean	[95% CI]	Mean	[95% CI]	Mean	[95% CI]	<i>p</i> -value
Functional performance tests							
30STS (reps)	10	[8.4, 11.1]	11	[10.2, 12.5]	1.6	[0.5, 2.6]	0.008*
TUG (s)	9.3	[8.4, 10.3]	8.6	[8.0, 9.3]	-0.7	[-1.3, -0.1]	0.032*
40MWT (m/s)	1.55	[1.47, 1.64]	1.63	[1.53, 1.73]	0.07	[0.03, 0.11]	0.003*
11-step SCT (s)	13.8	[12.3, 15.4]	13.9	[12.5, 15.4]	0.10	[-1.10, 1.31]	0.847
KOOS subscales							
Pain	72	[62, 82]	77	[70, 85]	5.3	[-3.2, 13.9]	0.218
Symptoms	80	[67, 92]	81	[73, 89]	1.2	[-7.8, 10.2]	0.778
ADL	76	[68, 84]	78	[72, 85]	2.2	[-5.0, 9.5]	0.528
Sport/Recreation	37	[21, 52]	45	[32, 58]	10.6	[-6.0, 27.1]	0.310
QOL	49	[43, 56]	48	[38, 58]	-1.4	[-5.0, 7.9]	0.642

30STS: 30 seconds chair-stand test; TUG: Timed Up and Go; 40MWT: 40m fast-paced walk test; 11-step SCT: 11-step stair-climb test; KOOS: Knee Injury and Osteoarthritis Outcome Score; ADL: activities of daily living; QOL: quality of life; 95% CI: 95% confidence interval. KOOS score 0 = extreme knee problems. Score 100 = no knee problems. *p < 0.05, Pre vs Post.

Functional performance

In the current study 8-10 weeks of BFR-walking exercise was accompanied by significant improvements in functional performance, assessed as 30STS (+16%), TUG (-8%) and 40MWT (+5%) in elderly individuals with knee OA. In contrast, stair walking performance remained unaltered following the period of training.

Improvements in functional performance have consistently been reported following BFR-walking exercise, which are in line with the current study data (14, 19, 24). Clarkson et al. (2017) demonstrated that 6 weeks of outdoor BFR-walking exercise (10-min sessions, 4km/h, 4 times a week) improved performance in 30 STS (+28 \pm 6%), TUG (-12 \pm 2%), 6MWT $(-9\pm1\%)$ and the modified Queen's College Step Test $(+80\pm11\%)$ in healthy elderly individuals (60-80 years). Likewise, Ozaki et al. (2011) demonstrated that 10 weeks of BFR treadmill walking exercise (20-min, 4.5km/h, 1.6°, 4 times a week) improved the performance in The Up & Go test (11%) and 30STS (21%) in healthy elderly individuals (57-73 years), while control subjects who performed regular walking without BFR experienced improvements only in the 30STS (8%).

The current study results combined with Clarkson et al. (2017) and Ozaki et al. (2011) indicate that BFRwalking exercise may represent an effective training modality to improve functional performance in elderly people, including individuals with knee OA.

Self-reported knee function

Unexpectedly, no changes in KOOS subscale scores were observed in the current study in response to 8-10 weeks of BFR-walking exercise, despite consistent improvements in functional performance. The lack of changes might be due to a ceiling effect of the KOOS scores. That is, a study by Steven-Lapsley et al. (2011) found a strong correlation between KOOS-Pain and KOOS-ADL scores in a group of patients undergoing total knee arthroplasty (39). KOOS was measured 2 weeks prior to surgery and 1-, 3- and 6-months post-surgery. KOOS-ADL and KOOS-Pain were continuously inter-related (r=0.766-0.826), indicating a strong relationship between pain intensity and ability to perform ADL. These data suggest that participants' perceived ability to perform given ADL-related activities may depend on perceived knee pain intensity. Therefore, given the relatively low levels of perceived knee pain observed at baseline, a ceiling effect on KOOS-ADL might have occurred in the present study. In line with this notion, the baseline KOOS subscale scores observed in the current study appeared to be higher compared with reference data reported previously in individuals with knee OA (40). Accordingly, the participants in this study might have difficulty improving their KOOS scores in general, due to already high levels of perceived knee function.

Study limitations

This study has a number of potential limitations. The absence of a non-exercising control group hinders determination of whether the BFR intervention was the direct cause of the concurrent improvements in functional performance. Furthermore, the small sample size gives rise to low statistical power in the pre-to-post comparisons. In addition, only 1 of 4 weekly training sessions were supervised and, during the period of COVID-19 restrictions, no on-site supervision was possible. Lastly, despite thorough education in applying the cuff proximally at the thigh, some participants experienced difficulties reproducing this procedure accurately, which could have influenced the effective occlusion pressure applied during BFR-walking.

Perspectives

There is a lack of evidence regarding the mechanisms behind increases in functional performance (including measures of muscle strength, hypertrophy and/ or neural drive) in response to BFR-walking exercise in patients with OA. Therefore, future studies examining the effect of BFR-walking on lower limb muscle strength, morphology and neuromuscular function in patients with knee OA seem highly relevant.

CONCLUSION

Nine out of 14 participants completed 8-10 weeks of BFR-walking exercise. The participants who completed the intervention period demonstrated high adherence to training and low-level knee pain during BFR-walking exercise. Furthermore, they demonstrated significant improvements in physical performance, albeit not improving self-reported knee function (KOOS). In contrast, 4 participants withdrew from the study due to intervention-related adverse events. Participants withdrawing from the study were characterized by a higher BMI, higher levels of perceived knee pain and low levels of physical performance, which may have negatively affected their ability to complete the exercise programme. Consequently, it becomes important to account for individual limitations when applying BFR-walking exercise to this population.

Regardless, the current data indicate that shortterm BFR-walking exercise can lead to significant improvements in physical performance, albeit not improving self-reported knee function (KOOS). Future randomized controlled trial (RCT) studies of more prolonged duration are required to provide stronger evidence on the effect of BFR-walking exercise in patients with knee OA.

ACKNOWLEDGEMENTS

We thank all study participants for the time they devoted to the study. Furthermore, we thank the staff at the local senior centre (Møllestien) in Aarhus (Denmark) for providing assistance during the recruitment period.

REFERENCES

- Hvidberg MF, Johnsen SP, Davidsen M, Ehlers L. A nationwide study of prevalence rates and characteristics of 199 chronic conditions in Denmark. Pharmacoecon Open 2020; 4: 361–380.
- Lane NE, Brandt K, Hawker G, Peeva E, Schreyer E, Tsuji W, et al. OARSI-FDA initiative: defining the disease state of osteoarthritis. Osteoarthritis Cartilage 2011; 19: 478–482.
- Alnahdi AH, Zeni JA, Snyder-Mackler L. Muscle impairments in patients with knee osteoarthritis. Sports Health 2012; 4: 284–292.
- Ikeda S, Tsumura H, Torisu T. Age-related quadricepsdominant muscle atrophy and incident radiographic knee osteoarthritis. J Orthop Sci 2005; 10: 121–126.
- Bennell K, Dobson F, Hinman R. Measures of physical performance assessments: Self-Paced Walk Test (SPWT), Stair Climb Test (SCT), Six-Minute Walk Test (6MWT), Chair Stand Test (CST), Timed Up & Go (TUG), Sock Test, Lift and Carry Test (LCT), and Car Task. Arthritis Care Res (Hoboken) 2011; 63 Suppl 11: S350–370.
- Skoffer B, Dalgas U, Mechlenburg I, Søballe K, Maribo T. Functional performance is associated with both knee extensor and flexor muscle strength in patients scheduled for total knee arthroplasty: a cross-sectional study. J Rehabil Med 2015; 47: 454–459.
- Araujo IL, Castro MC, Daltro C, Matos MA. Quality of life and functional independence in patients with osteoarthritis of the knee. Knee Surg Relat Res 2016; 28: 219–224.
- Huang KH, Hsieh RL, Lee WC. Pain, physical function, and health in patients with knee osteoarthritis. Rehabil Nurs 2017; 42: 235–241.
- Sutbeyaz ST, Sezer N, Koseoglu BF, Ibrahimoglu F, Tekin D. Influence of knee osteoarthritis on exercise capacity and quality of life in obese adults. Obesity (Silver Spring) 2007; 15: 2071–2076.
- Ferraz RB, Gualano B, Rodrigues R, Kurimori CO, Fuller R, Lima FR, et al. Benefits of resistance training with blood flow restriction in knee osteoarthritis. Med Sci Sports Exerc 2018; 50: 897–905.
- Li Y, Su Y, Chen S, Zhang Y, Zhang Z, Liu C, et al. The effects of resistance exercise in patients with knee osteoarthritis: a systematic review and meta-analysis. Clin Rehabil 2016; 30: 947–959.
- Skoffer B, Maribo T, Mechlenburg I, Hansen PM, Søballe K, Dalgas U. Efficacy of preoperative progressive resistance training on postoperative outcomes in patients undergoing total knee arthroplasty. Arthritis Care Res (Hoboken) 2016; 68: 1239–1251.
- Jan MH, Lin JJ, Liau JJ, Lin YF, Lin DH. Investigation of clinical effects of high- and low-resistance training for patients with knee osteoarthritis: a randomized controlled trial. Phys Ther 2008; 88: 427–436.
- 14. Abe T, Sakamaki M, Fujita S, Ozaki H, Sugaya M, Sato Y, et al. Effects of low-intensity walk training with restricted leg blood flow on muscle strength and aerobic capacity in older adults. J Geriatr Phys Ther 2010; 33: 34–40.
- 15. Bryk FF, Dos Reis AC, Fingerhut D, Araujo T, Schutzer M, Cury Rde P, et al. Exercises with partial vascular occlusion

in patients with knee osteoarthritis: a randomized clinical trial. Knee Surg Sports Traumatol Arthrosc 2016; 24: 1580–1586.

- Clarkson MJ, May AK, Warmington SA. Chronic blood flow restriction exercise improves objective physical function: a systematic review. Front Physiol 2019; 10: 1058.
- 17. Grønfeldt BM, Lindberg Nielsen J, Mieritz RM, Lund H, Aagaard P. Effect of blood-flow restricted vs heavy-load strength training on muscle strength: Systematic review and meta-analysis. Scand J Med Sci Sports 2020; 30: 837–848.
- Hughes L, Paton B, Rosenblatt B, Gissane C, Patterson SD. Blood flow restriction training in clinical musculoskeletal rehabilitation: a systematic review and meta-analysis. Br J Sports Med 2017; 51: 1003–1011.
- Ozaki H, Sakamaki M, Yasuda T, Fujita S, Ogasawara R, Sugaya M, et al. Increases in thigh muscle volume and strength by walk training with leg blood flow reduction in older participants. J Gerontol A Biol Sci Med Sci 2011; 66: 257–263.
- Wernbom M, Aagaard P. Muscle fibre activation and fatigue with low-load blood flow restricted resistance exercise-An integrative physiology review. Acta Physiol (Oxf) 2020; 228: e13302.
- Haddock B, Hansen SK, Lindberg U, Nielsen JL, Frandsen U, Aagaard P, et al. Physiological responses of human skeletal muscle to acute blood flow restricted exercise assessed by multimodal MRI. J Appl Physiol (1985) 2020; 129: 748–759.
- Takada S, Okita K, Suga T, Omokawa M, Kadoguchi T, Sato T, et al. Low-intensity exercise can increase muscle mass and strength proportionally to enhanced metabolic stress under ischemic conditions. J Appl Physiol (1985) 2012; 113: 199–205.
- Nielsen JL, Aagaard P, Bech RD, Nygaard T, Hvid LG, Wernbom M, et al. Proliferation of myogenic stem cells in human skeletal muscle in response to low-load resistance training with blood flow restriction. J Physiol 2012; 590: 4351–4361.
- Clarkson MJ, Conway L, Warmington SA. Blood flow restriction walking and physical function in older adults: a randomized control trial. J Sci Med Sport 2017; 20: 1041–1046.
- Kacin A, Rosenblatt B, Grapar Zargi T, Biswas A. Safety considerations with blood flow restricted resistance training. Annales Kinesiologiae 2015; 6: 3–26.
- Karabulut M, Sherk VD, Bemben DA, Bemben MG. Inflammation marker, damage marker and anabolic hormone responses to resistance training with vascular restriction in older males. Clin Physiol Funct Imaging 2013; 33: 393–399.
- Minniti MC, Statkevich AP, Kelly RL, Rigsby VP, Exline MM, Rhon DI, et al. The safety of blood flow restriction training as a therapeutic intervention for patients with musculoskeletal disorders: a systematic review. Am J Sports Med 2020; 48: 1773–1785.
- Pope ZK, Willardson JM, Schoenfeld BJ. Exercise and blood flow restriction. J Strength Cond Res 2013; 27: 2914–2926.
- 29. Slysz J, Stultz J, Burr JF. The efficacy of blood flow restricted exercise: a systematic review & meta-analysis. J Sci Med Sport 2016; 19: 669–675.
- Petersson N, Jørgensen S, Kjeldsen T, Aagaard P, Mechlenburg I. Okklusionsgangtræning som genoptræning til en patient med kronisk knæartrose. [Blood-flow restricted walking exercise as rehabilitation for a patient with chronic knee osteoarthritis]. Ugeskr Laeger 2020; 182 (41).
- Kilgas MA, McDaniel J, Stavres J, Pollock BS, Singer TJ, Elmer SJ. Limb blood flow and tissue perfusion during exercise with blood flow restriction. Eur J Appl Physiol 2019; 119: 377–387.
- Patterson SD, Hughes L, Warmington S, Burr J, Scott BR, Owens J, et al. Blood flow restriction exercise position stand: considerations of methodology, application, and safety. Front Physiol 2019; 10: 533.
- 33. Jørgensen SL, Bohn MB, Aagaard P, Mechlenburg I. Efficacy of low-load blood flow restricted resistance exercise in patients with knee osteoarthritis scheduled for total

Journal of Rehabilitation Medicine

knee replacement (EXKnee): protocol for a multicentre randomised controlled trial. BMJ Open 2020; 10: e034376.

- Dobson F, Hinman RS, Hall M, Terwee CB, Roos EM, Bennell KL. Measurement properties of performance-based measures to assess physical function in hip and knee osteoarthritis: a systematic review. Osteoarthritis Cartilage 2012; 20: 1548–1562.
- 35. Tolk JJ, Janssen RPA, Prinsen CAC, Latijnhouwers DAJM, van der Steen MC, Bierma-Zeinstra SMA, et al. The OARSI core set of performance-based measures for knee osteoarthritis is reliable but not valid and responsive. Knee Surg Sports Traumatol Arthrosc 2019; 27: 2898–2909.
- 36. Collins NJ, Misra D, Felson DT, Crossley KM, Roos EM. Measures of knee function: International Knee Documentation Committee (IKDC) Subjective Knee Evaluation Form, Knee Injury and Osteoarthritis Outcome Score (KOOS), Knee Injury and Osteoarthritis Outcome Score Physical Function Short Form (KOOS-PS), Knee Outcome Survey Activities of Daily Living Scale (KOS-ADL), Lysholm Knee Scoring Scale, Oxford Knee Score (OKS), Western Ontario and McMaster

Universities Osteoarthritis Index (WOMAC), Activity Rating Scale (ARS), and Tegner Activity Score (TAS). Arthritis Care Res (Hoboken) 2011; 63 Suppl 11: S208–228.

- 37. Roos EM, Lohmander LS. The Knee injury and Osteoarthritis Outcome Score (KOOS): from joint injury to osteoarthritis. Health Qual Life Outcomes 2003; 1: 64.
- Gomes-Neto M, Araujo AD, Junqueira ID, Oliveira D, Brasileiro A, Arcanjo FL. Comparative study of functional capacity and quality of life among obese and non-obese elderly people with knee osteoarthritis. Rev Bras Reumatol Engl Ed 2016; 56: 126–130.
- 39. Stevens-Lapsley JE, Schenkman ML, Dayton MR. Comparison of self-reported knee injury and osteoarthritis outcome score to performance measures in patients after total knee arthroplasty. Pm r 2011; 3: 541–549; quiz 9.
- Collins NJ, Prinsen CA, Christensen R, Bartels EM, Terwee CB, Roos EM. Knee Injury and Osteoarthritis Outcome Score (KOOS): systematic review and meta-analysis of measurement properties. Osteoarthritis Cartilage 2016; 24: 1317–1329.