

Walking with Leg Blood Flow Restriction: Wide-Rigid Cuffs vs. Narrow-Elastic Bands

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Conflict of interest statement

The authors declare a potential conflict of interest and state it below

A conflict of interest was declared by Sten Stray-Gundersen, who is presently employed by BStrongTm, Park City, Utah. For the remaining authors, none were declared

Author contribution statement

All authors contributed to the generation of study idea, data collection and analyses, and drafting of the manuscript. All authors approved the final version of the manuscript.

Keywords

Exercise, Cardiovascular disease, Exercise Training, Blood flow restricted exercise, Physical exercise, hemodynamic stress

Abstract

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Background: Blood flow restriction (BFR) training is becoming a popular form of exercise. Walking exercise in combination with pressurized wide-rigid (WR) cuffs elicits higher cardiac workload and a vascular dysfunction due presumably to reperfusion injury to the endothelium. In contrast, narrow-elastic (NE) BFR bands may elicit different hemodynamic effects. Therefore, we compared the acute cardiovascular responses to two distinct forms of BFR training during light-intensity exercise. Methods and Results: 15 young healthy participants (M=9, F=6) performed 5 bouts of 2-minute walking intervals at 0.9 m/s with a 1-minute rest and deflation period with either WR, NE, or no bands placed on upper thighs. Cuff pressure was inflated to 160 mmHg in WR cuffs and 300 mmHg in NE bands while no cuffs were used for the control. Increases in heart rate and arterial blood pressure were greater ($p<0.05$) in the WR than the NE and control conditions. Double product increased to a greater extent in the WR than in the NE and control conditions. Increases in perceived exertion and blood lactate concentration were greater ($p<0.05$) in the WR compared with the NE and control conditions ($p<0.05$), while no differences emerged between the NE and control conditions. There were no changes in arterial stiffness or brachial artery flow-mediated dilation after all three trials. Conclusion: Use of wide-rigid BFR cuffs resulted in a marked increase in blood pressure and myocardial oxygen demand compared with narrow-elastic BFR bands, suggesting that narrow-elastic bands present a safer alternative for at-risk populations to perform BFR exercise. Clinical Trial Registration: This study was registered in the Clinicaltrials.gov (NCT03540147).

Contribution to the field

Attached please find our manuscript entitled "Walking with leg blood flow restriction: Wide-rigid cuffs vs. narrow-elastic bands" submitted to the Journal of the American Heart Association. This manuscript represents results of original work that have not been published elsewhere (except as an abstract in conference proceedings). This manuscript has not and will not be submitted for publication elsewhere until a decision is made regarding its acceptability for publication in the Journal of the American Heart Association. If accepted for publication, it will not be published elsewhere. Over the past two decades, blood flow restriction (BFR) training has increased in popularity among athletes and has been increasingly prescribed to older patients with cardiovascular diseases. Concern has been raised over the use of BFR in at-risk populations. One such complication could be an augmentation of the exercise pressor reflex, which is exaggerated in certain at-risk populations. We found that use of wide-rigid BFR cuffs resulted in a marked increase in blood pressure and myocardial oxygen demand compared with narrow-elastic BFR bands, suggesting that narrow-elastic bands present a safer alternative for at-risk populations to perform BFR exercise.

Ethics statements

Studies involving animal subjects

Generated Statement: No animal studies are presented in this manuscript.

Studies involving human subjects

Generated Statement: The studies involving human participants were reviewed and approved by IRB at the University of Texas at Austin. The patients/participants provided their written informed consent to participate in this study.

Inclusion of identifiable human data

Generated Statement: No potentially identifiable human images or data is presented in this study.

In review

Data availability statement

Generated Statement: The raw data supporting the conclusions of this article will be made available by the authors, without undue reservation.

In review

1 Walking with Leg Blood Flow Restriction: Wide-Rigid Cuffs vs. 2 Narrow-Elastic Bands

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9 **Keywords:** exercise; cardiovascular Disease; exercise physiology; exercise training; physical
10 exercise; hemodynamic stress

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12 exercise in combination with pressurized wide-rigid (WR) cuffs elicits higher cardiac workload and a
13 vascular dysfunction due presumably to reperfusion injury to the endothelium. In contrast, narrow-
14 elastic (NE) BFR bands may elicit different hemodynamic effects. Therefore, we compared the acute
15 cardiovascular responses to two distinct forms of BFR training during light-intensity exercise.

16 **Methods and Results:** 15 young healthy participants (M=9, F=6) performed 5 bouts of 2-minute
17 walking intervals at 0.9 m/s with a 1-minute rest and deflation period with either WR, NE, or no
18 bands placed on upper thighs. Cuff pressure was inflated to 160 mmHg in WR cuffs and 300 mmHg
19 in NE bands while no cuffs were used for the control. Increases in heart rate and arterial blood
20 pressure were greater ($p<0.05$) in the WR than the NE and control conditions. Double product
21 increased to a greater extent in the WR than in the NE and control conditions. Increases in perceived
22 exertion and blood lactate concentration were greater ($p<0.05$) in the WR compared with the NE and
23 control conditions ($p<0.05$), while no differences emerged between the NE and control conditions.
24 There were no changes in arterial stiffness or brachial artery flow-mediated dilation after all three
25 trials. **Conclusion:** Use of wide-rigid BFR cuffs resulted in a marked increase in blood pressure and
26 myocardial oxygen demand compared with narrow-elastic BFR bands, suggesting that narrow-elastic
27 bands present a safer alternative for at-risk populations to perform BFR exercise. **Clinical Trial**
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29 1 Introduction

30 Over the past two decades, blood flow restriction (BFR) training has increased in popularity
31 among athletes, researchers, and physical therapists (1). During this form of training, users place
32 pressurized cuffs/bands or non-pressurized straps/wraps on the most proximal portion of the limb in
33 order to restrict venous blood flow while maintaining varying degrees of arterial inflow (2). The
34 restriction of venous blood flow while performing light-weight exercises leads to venous pooling and
35 local metabolic changes that together stimulate systemic adaptations similar to those achieved with
36 heavy exercise (3, 4). Since BFR used in combination with low-intensity walking exercise can confer
37 significant improvements in muscle strength and hypertrophy (3, 5), there is great potential for use
38 with clinical populations for fitness and rehabilitation.

39 Concern has been raised over the use of BFR in at-risk populations (e.g., hypertensive, obese,
40 atherosclerotic) due to the potential for deep vein thrombosis, rhabdomyolysis, pulmonary emboli (6,

7) and other serious complications associated with occluding arterial flow and performing skeletal muscle contractions. One such complication could be an augmentation of the exercise pressor reflex, which is exaggerated in certain at-risk populations (8), and is normally elicited during exercise by the stimulation of group III and IV afferents (local mechano- and metaboreceptors), resulting in a sympathetically-mediated elevation in blood pressure and heart rate. Since BFR training leads to an accumulation of metabolites and exerts high pressures on blood vessels and contracting muscle tissue, it seems likely to elicit an exaggerated blood pressure response (9). Indeed, wide-rigid BFR cuffs can cause painful compression of tissues, increases in systemic vascular resistance, acute vascular dysfunction, and increased myocardial demand even at low exercise intensities (10, 11).

However, a multitude of Japanese athletes, seniors, clinicians, and trainers have been using BFR in the form of Kaatsu for over 30 years with an extremely low incidence of serious complications (6, 7). The more recent findings and resulting concerns may be due to a shift from the original narrow-elastic design present in the Kaatsu bands to wide-rigid nylon cuffs adapted from surgical tourniquets and blood pressure cuffs. The wide-rigid cuffs are easily available, but have the potential to inhibit the expansion of muscle upon increased blood flow accompanying exercise and muscle contraction while the narrow-elastic bands do not prevent the expansion.

To elucidate the potentially differing effects of these two distinct forms of BFR, we assessed the acute hemodynamic responses of fifteen young healthy individuals during low-intensity walking exercise while using wide-rigid cuffs or narrow-elastic bands. We hypothesized that the wide-rigid cuffs would elicit greater pressor responses and myocardial oxygen demand compared with the narrow-elastic bands during light-intensity aerobic exercise. Additionally, we hypothesized that systemic endothelial function and arterial stiffness will not be affected by an acute bout of BFR walking exercise, regardless of cuff type. If discovered that the narrow-elastic bands do not elicit the same heightened blood pressure responses to BFR, they may present a safer alternative for at-risk populations predisposed to exhibit exaggerated hemodynamic responses during exercise. We studied young healthy adults as they are the biggest users of BFR and to first determine whether the hemodynamic responses observed were within safe ranges before extending the studies to more vulnerable populations.

BFR using wide-rigid cuffs may induce an ischemia-reperfusion injury to the distal vessels upon the release of ischemia (10). One of the hallmark features of the ischemia-reperfusion injury is endothelial dysfunction leading to arterial stiffening. These vascular changes induced by the BFR exercise could be unfavorable or even detrimental to those with compromised cardiovascular conditions. Indeed our previous investigation (10) found a reduction in popliteal FMD after performing submaximal walking bouts with wide-rigid cuffs placed on thighs. Accordingly, we determined whether this effect was systemic in nature or localized to the occluded artery by assessing brachial endothelial function after the BFR exercise with cuffs placed on thighs.

2 Methods

2.1 Participants

A total of 15 young healthy sedentary and recreationally active adults (9 males and 6 females) between the ages of 18 and 35 years participated in this study. Exclusion criteria for participation, assessed via medical history questionnaire, included (1) uncontrolled hypertension; (2) smoking within the last 6 months; (3) a history of heart disease, kidney disease, peripheral artery disease, and other known cardiovascular issues; (4) obesity as defined by a body mass index (BMI) >30 kg/m²; (5) a history of diabetes or other metabolic dysfunction; (6) major operations within the last 6 months; (7) advised to avoid exercise by a physician; (8) part of a vulnerable population (unable to consent, pregnant women, osteoporotic, etc.): or (9) currently performing BFR training. All participants

87 submitted their written informed consent prior to participation. The Institutional Review Board
88 reviewed and approved this study.

89

90 2.2 Procedures

91 Participants visited the laboratory on 2 separate occasions for 2 hours per visit. During the first visit,
92 anthropometric measures of height, body weight, and body fatness were taken. Body fatness was
93 estimated using the 7-site skinfold technique with Lange skinfold calipers (Beta Technology, Santa
94 Cruz, CA). Participants fasted for at least 8 hours, did not consume alcohol or caffeine for 12 hours,
95 and abstained from strenuous exercise for 24 hours prior to each experimental session.

96 After 20 minutes of supine rest in a quiet, temperature-controlled room (23–27°C), baseline
97 measurements, including heart rate, blood pressure using an automated sphygmomanometer, arterial
98 stiffness using a pulse-wave velocity index, and brachial endothelial function via flow-mediated
99 dilation, were conducted. After undergoing baseline measurements, each participant performed one of
100 three randomly-assigned walking exercise conditions; walking with pressurized wide-rigid cuffs (WR),
101 walking with pressurized narrow-elastic bands (NE), or walking without cuffs/bands (control).
102 Cuffs/bands were placed on both legs and subsequently inflated when performing one of the BFR
103 conditions while no cuffs were used when performing the control. In men, visits were separated by at
104 least three days. In women, visits were scheduled ~ 1 month apart during the early follicular phase of
105 the menstrual cycle approximately 1-5 days following the start of menstruation. All participants
106 performed the three conditions in a randomized order on three separate days. In addition to taking
107 plasma lactate samples immediately before and after exercise, we assessed RPE before, mid-way
108 through, and after exercise. During the exercise, we recorded beat-by-beat blood pressure and heart
109 rate continuously via finger plethysmography. Once the participant completed the exercise, measures
110 of vascular functions were repeated within a 15-minute period and then one hour after the completion
111 of the exercise.

112

113 2.3 Measurements

114 Heart rate at rest, brachial blood pressure, and arterial stiffness as assessed by cardio-ankle
115 vascular index (CAVI) were measured in the supine position using the automated vascular screen
116 device (VaSera, Fakuda Denshi, Tokyo, Japan) as previously described (12).

117 Flow-mediated dilation (FMD), a measure of vascular endothelium-dependent vasodilation was
118 assessed using a semi-automated diagnostic ultrasound system with a semi-automated probe, which
119 self-adjusts to provide clear images of the intimal layer for baseline artery diameter measurements (EF-
120 38G, UNEX corporation, Nagoya, Japan) (13). While participants rested in the supine position, a
121 pneumatic cuff was placed on the right forearm. Then cross-sectional and longitudinal images of the
122 brachial artery were acquired 6-8 cm proximal to the cuff. In order to occlude blood flow, the cuff was
123 subsequently inflated to 50 mmHg above resting systolic blood pressure for a period of five minutes.
124 Upon cuff deflation, blood flow velocity and artery diameter were measured for an additional two
125 minutes.

126 Blood lactate concentration was measured immediately before and between 90 and 120 seconds
127 after the walking protocol. Using disposable lancets, we punctured the finger-tip and collected a blood
128 droplet on a disposable lactate test-strip. We did not warm the fingers prior to the finger-prick as it was
129 not necessary for this population. All blood samples were analyzed using a portable lactometer
130 (LactatePro, Arkray, Kyoto, Japan).

131 Ratings of perceived exertion were assessed before, during, and after the walking protocol.
132 Participants were familiarized with the scale prior to the beginning of the test and asked to score their
133 perceived exertion using the original Borg scale.

134 In order to record hemodynamics during the walking protocol, beat-to-beat arterial blood
135 pressure waveforms were continuously measured via finger plethysmography (Portapres Model 2,
136 TNO TPD Biomedical Instruments, Netherlands) placed on the middle finger of the left hand of each
137 participant. Following standard procedure in order to control for potential changes in hydrostatic
138 pressure due to variable hand position, participants were instructed to keep the left hand at heart level
139 during the entirety of the exercise session. The participant's right hand was free to move in a normal
140 walking fashion or to stabilize themselves during a trip or fall. Heart rate was calculated from the finger
141 blood pressure waveform using the validated model-flow method (BeatScope 1.0 software, TNO TPD
142 Biomedical Instrumentation, Amsterdam, The Netherlands). Double product, an index of myocardial
143 oxygen demand, was calculated by systolic blood pressure \times heart rate. The hemodynamic values
144 represent the average values during the 2-minute walking bout, excluding the 1-minute rest interval.

145 146 2.4 Exercise Protocol

147 The walking exercise test consisted of five bouts of 2-minute walking intervals at 0.9 m/s with
148 a 1-minute rest and deflation period between each bout with either wide-rigid cuffs, narrow-elastic
149 bands, or no cuffs placed on the upper thighs (10, 11, 14). The chosen treadmill speed has been used
150 in previous investigations in our laboratory as a means to evoke a submaximal effort, and is a typical
151 speed used during cardiac rehabilitation programs (10, 11, 14). We used two commercially-available
152 cuffs as representatives of wide-rigid cuffs and narrow-elastic bands. For the wide-rigid cuff condition,
153 we used wide rapid-inflation pneumatic tourniquets (Hokanson, CC17, Bellevue, WA; 18 cm wide \times
154 108 cm long). We used a one-size-fits-all thigh-cuff typically used by people performing lower limb
155 BFR, and did not observe any differences in responses between smaller and larger participants. For the
156 narrow-elastic band condition, we used pneumatically-controlled BFR leg bands (BStrong, Park City,
157 UT; 5 cm wide \times 50 cm long). Following previous protocols (10, 14), and in order to familiarize the
158 participant with the wide-rigid cuff, we initially inflated the cuff to 120 mmHg for 30 seconds, released
159 it for 10 seconds, re-inflated to 140 mmHg for 30 seconds, released for 10 seconds, and then re-inflated
160 to the final pressure of 160 mmHg. Standing baseline heart rate and blood pressure was recorded via
161 finger plethysmography for 1-minute before beginning the walking exercise. For the narrow-elastic
162 condition, we gradually inflated the bands to 300 mmHg, which is the recommended and commonly-
163 used pressure for leg BFR according to the company supplying the equipment. These same standard
164 pressures were used for all individuals for comparative purposes. Once the desired pressure was
165 reached, the participants began the walking exercise. After completion of each 2-minute bout, we
166 rapidly deflated both cuffs for 1-minute before the next bout. After the fifth bout, we continued
167 recording blood pressure and heart rate for one additional minute. In the control session, participants
168 performed the same exercise protocol without the application or inflation of either cuff.

169 170 2.5 Statistical Analyses

171 Parametric statistics were used as the data were normally distributed as determined by a
172 Levene's test. Since baseline measures for systemic hemodynamics were not different, one-way
173 repeated measures ANOVA was used to identify significant effects across the three conditions. For
174 ratings of perceived exertion, blood lactate, CAVI, and flow-mediated dilation, two-way repeated
175 measures ANOVA was used. After determining whether significant main effects or interactions
176 ($p < 0.05$) were present, we ran post-hoc multiple comparison t-tests ($p < 0.05$) using a Bonferroni
177 correction to identify the significant differences between conditions. Data are presented as
178 means \pm SEM unless stated otherwise.

179 **3 Results**

180 Selected participant characteristics are presented in Table 1. Participants were young, healthy, and
181 exhibited normal body weight and composition. Absolute values for selected hemodynamic variables
182 before and during each 2-minute walking bout are presented in Table 2. Changes in arterial blood
183 pressure from baseline during each 2-minute walking bout are presented in Figure 1. At baseline, no
184 differences existed in any of the variables between the three conditions. Increases in blood pressure
185 were greater ($p < 0.05$) in the wide-rigid cuff condition than the narrow-elastic band and control
186 conditions while increases in systolic and mean arterial blood pressure were greater in the control
187 compared with the narrow-elastic condition ($p < 0.05$). As presented in Figure 2, increases in double
188 product were greater ($p < 0.05$) in the wide-rigid cuff condition than the narrow-elastic band and
189 control conditions and increases were greater in the control compared with the narrow-elastic
190 condition ($p < 0.05$). Increases in heart rate were greater ($p < 0.05$) in the wide-rigid condition than the
191 narrow-elastic and control conditions and were not different between the narrow-elastic condition
192 and control ($p > 0.05$). Blood lactate concentrations measured before and immediately after walking
193 are presented in Figure 3. Increases in blood lactate concentrations were greater ($p < 0.05$) in the wide-
194 rigid cuff condition than the narrow-elastic band and control conditions. As shown in Figure 4,
195 ratings of perceived exertion were greater ($p < 0.05$) in the wide-rigid condition than the control
196 condition immediately post-exercise. Cardio-ankle vascular index (CAVI) and flow-mediated
197 dilation (FMD) did not change across all three conditions (Figure 5).

198

199

200 4 Discussions

201 The present study aimed to evaluate hemodynamic responses between two distinct forms of
202 BFR training commonly used by trainers, physical therapists, and researchers. In agreement with
203 previous investigations (10, 11), the use of wide-rigid BFR cuffs elicited markedly increased blood
204 pressure responses and heightened myocardial oxygen demands during light intensity walking
205 compared with the narrow-elastic bands and control conditions. In contrast, the use of narrow-elastic
206 bands did not elicit increased hemodynamic responses compared to control, suggesting that narrow-
207 elastic BFR does not appear to pose additional risk to users than light-intensity walking without BFR.
208 In fact, systolic blood pressure, mean arterial blood pressure, and double product values were greater
209 in the control condition compared with the narrow-elastic condition. None of the conditions induced
210 acute measurable changes in cardio-ankle vascular indices or flow-mediated dilation, suggesting that
211 these forms of BFR do not promote systemic vascular dysfunction or arterial stiffening. These
212 findings are novel and suggest that narrow-elastic BFR may present a safe option for at-risk
213 populations to perform BFR as a mode of exercise and rehabilitation.

214 The mechanisms underlying the differing hemodynamic and metabolic responses to the two
215 forms of BFR exercise remain elusive and beyond the scope of this investigation as these variables
216 were not measured in the present study due to technical issues. However, it is clear that the width and
217 material of the cuff have profound effects on systemic hemodynamics. This is likely due to varying
218 degrees of arterial occlusion and compression of muscle tissue leading to variable increases in blood
219 pressure, systemic vascular resistance, and local mechanoreceptor stimulation (2, 9). In particular, the
220 use of wide-rigid cuffs results in highly compressive forces over a large area of the limb, inhibiting
221 the muscle from expanding with an increase in blood flow accompanying exercise. In contrast, a
222 narrow-elastic design appears to minimize how much working muscle is compressed during repeated
223 muscle contractions, allowing the muscle to swell upon increased blood flow. This is evidenced by
224 the slight drop in diastolic blood pressure observed during the control and narrow-elastic conditions.

225 It appears that narrow-elastic BFR systems provide a wide range of pressures in which one can avoid
226 arterial occlusion, while sufficiently restricting venous outflow to create a disturbance of homeostasis
227 in working muscle. With wide-rigid BFR systems, due to the rigid outer material that cannot expand,
228 the cuff is not able to accommodate the increase in cross-sectional area when muscle contracts and
229 increases its cross-sectional area—pressures spike in the tissues contained by the cuff, arteries
230 become occluded, and veins remain closed. Thus, the wide-rigid BFR system stops functioning
231 correctly and complications like peripheral nerve injury, rhabdomyolysis, and deep venous thrombi
232 (DVT's) become more likely.

233 The elastic nature of bands enables the distal portion of the muscle to force blood past the
234 intermittent venous blockade, minimizing the degree of pain, arterial occlusion, and mechanical
235 compression of tissues. Furthermore, since BFR can be equally effective at inducing hypertrophy and
236 strength gains at 40% and 90% arterial occlusion with the use of wide-rigid cuffs (15), the percent
237 arterial occlusion does not appear to be of primary concern for an effective BFR session. This is of
238 considerable importance given that an increase in systemic vascular resistance, which is elevated
239 when occluding any percentage of arterial inflow, leads to increases in blood pressure and heart rate
240 (9). In contrast, during aerobic exercise of varying intensities, systemic vascular resistance decreases
241 slightly due to a vasodilatory response in the working muscle to exercise (16). Therefore, the
242 observed increases in systemic vascular resistance during BFR (10, 17) are likely a result of the
243 mechanical constriction of muscle and other tissues beneath the cuff that functionally inhibits tissue
244 expansion and reduces the effect of the local vasodilatory response elicited during aerobic exercise.
245 When using narrow-elastic bands, this mechanically-mediated rise in systemic vascular resistance
246 appears to be absent as muscle contractions are able to intermittently pump blood past the venous
247 impediment and induce peripheral vasodilation. This is primarily evidenced by the considerable
248 increase in diastolic pressure while using the wide-rigid cuffs compared with the narrow-elastic and
249 control conditions. This suggests that the exercise pressor reflex only becomes exaggerated when
250 using wide-rigid cuffs at a commonly-used pressure of 160 mmHg, prompting the need for
251 individualized pressures and/or use of narrow-elastic bands with at-risk populations.

252 The lower systolic and mean blood pressure in the narrow elastic band condition compared
253 with the control condition that we observed in the present study is perplexing. In the present study,
254 heart rate was not different between the narrow-elastic band and the control conditions. Since
255 systolic blood pressure is driven by changes in stroke volume (18), it is possible that the narrow-
256 elastic band somehow facilitated venous blood pooling, which in turn reduced venous return, preload,
257 stroke volume resulting in lower systolic blood pressure. Indeed the previous investigation (10) found
258 that increases in stroke volume were attenuated in the BFR condition compared with the control.

259 In addition to exaggerated hemodynamic responses, the wide-rigid cuffs elicited significantly
260 higher ratings of perceived exertion and increases in plasma lactate concentrations. This suggests a
261 greater accumulation of metabolites in the muscle during the wide-rigid condition than the narrow-
262 elastic and control conditions given the same absolute workload. Although speculative, this could be
263 due to a greater degree of arterial impediment leading to more anaerobic metabolism, and more
264 venous blood pooling as the skeletal muscle pump is unable to push metabolite-rich blood past the
265 occluding cuff. This may be useful for healthy individuals performing low loads in a highly
266 controlled environment such as a physical therapist clinic. However, it also poses the risk of an
267 augmentation of the exercise pressor reflex, leading to unnecessary increases in blood pressure to
268 achieve the desired BFR stimulus.

269 In agreement with a past investigation (19), participants frequently complained of pain from
270 the compression of the cuff when using the wide-rigid cuffs while there were no complaints when
271 using the narrow-elastic bands. This may have confounded the ratings of perceived exertion seen in
272 the wide-rigid condition as pain can augment relative measures of effort during certain types of
273 exercise (20). However, pain also stimulates sympathetic activity, which could lead to an even

274 greater augmentation of the exercise pressor reflex (9). Therefore, more investigation into the use of
275 narrow-elastic bands during intense exercise, and the subsequent increase in muscle pain is warranted
276 to determine the mechanism responsible for the differences observed.

277 In a previous study (10), walking in combination with wide-rigid cuffs acutely decreased
278 endothelial function as assessed via flow-mediated dilation of the popliteal artery. To determine
279 whether the acute endothelial dysfunction previously observed was localized to the vasculature that
280 was exposed to ischemia and reperfusion or a sign of systemic endothelial dysfunction, we measured
281 brachial artery diameter changes in response to post-occlusive reactive hyperemia. The flow-
282 mediated dilation values remained unchanged across all conditions and time points, suggesting that
283 the acute decrease in popliteal flow-mediated dilation previously observed was likely a consequence
284 of local ischemia-reperfusion injury to the vascular endothelium distal to the cuff, and not a drop in
285 systemic endothelial function.

286 There were several limitations to this study. As the purpose of this study was to acutely
287 determine the relative safety among different forms of BFR in young healthy individuals, there was
288 no measure of the degree of efficacy between cuffs. Additionally, the responses of young healthy
289 individuals do not necessarily translate to more vulnerable populations. Moreover, since this study
290 only assessed acute responses to walking exercise with BFR, we cannot definitely say whether these
291 responses would be the same for long-term aerobic or resistance training. Clearly, there is a need for
292 further research investigating the cardiovascular effects of various forms of BFR and exercise
293 protocols.

294 There are several opportunities to further elucidate the mechanisms for the observed responses
295 in the present study as well as the use of BFR in various populations and protocols. Firstly, as the
296 present study assessed differences between wide-rigid and narrow-elastic cuffs, further research
297 showing the potential differences between narrow-rigid and wide-elastic cuffs are needed to further
298 elucidate the cause of the increased pressor response. Although hemodynamic responses to various
299 forms of BFR have been investigated at rest (21), the same measures need to be conducted during
300 exercise. In addition, there is a need for more research on BFR as a rehabilitation tool for hypertensive
301 patients. Although several researchers have found evidence of an attenuation of hypertension after a
302 BFR training program (17, 22-24), more investigation into the precise mechanism as well as the use of
303 a variety of training protocols is necessary. There is a spectrum of BFR equipment that can elicit the
304 desired stimulus at various pressures and intensities (25), so individually determining the optimal type
305 of equipment and exercise protocols could provide opportunities for a wide range of the population to
306 use BFR. In conclusion, the main finding in the present study is that the use of wide-rigid BFR cuffs
307 elicited markedly increased pressor responses and a heightened myocardial oxygen demand during
308 low-intensity walking compared with the narrow-elastic bands or control. It appears that an
309 exaggerated blood pressure response should be expected when using wide-rigid BFR cuffs that increase
310 systemic vascular resistance by occluding arterial inflow, compressing tissues, and reducing the ability
311 of the skeletal muscle pump to function. Therefore, we conclude that wide-rigid cuffs may only be safe
312 within a narrow window of pressures and should be conducted in a setting in which continuous
313 hemodynamics are monitored. In contrast, the narrow-elastic bands do not seem to elicit an augmented
314 exercise pressor response compared to control. These findings suggest that at-risk populations can
315 perform BFR without fear of overt cardiovascular risk. By nature of its width and material, it is difficult
316 to minimize the risks associated with wide-rigid cuffs when occluding any amount of arterial blood
317 flow, and as such, it should be prescribed carefully.

318 **5 Conflict of Interest**

319 *A potential conflict of interest was declared by Sten Stray-Gundersen, whose family members are*
320 *employed by BStrong™, Park City, Utah. For the remaining authors, none were declared. The results*

321 *of the present study do not constitute endorsement by ACSM. The results of the study are presented*
322 *clearly, honestly, and without fabrication, falsification, or inappropriate data manipulation.*

323 **6 Author Contributions**

324 SG was responsible for data collection, data analysis, authoring of the manuscript, and generation of
325 figures and tables. SW aided in data collection, data acquisition, and data analysis. HT was
326 responsible for study design, overseeing the entirety of the project, editing and reviewing the
327 manuscript, and provided final formatting of the document.

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In review

Figure 1 JPEG

○ Control

■ Narrow Elastic

▲ Wide Rigid

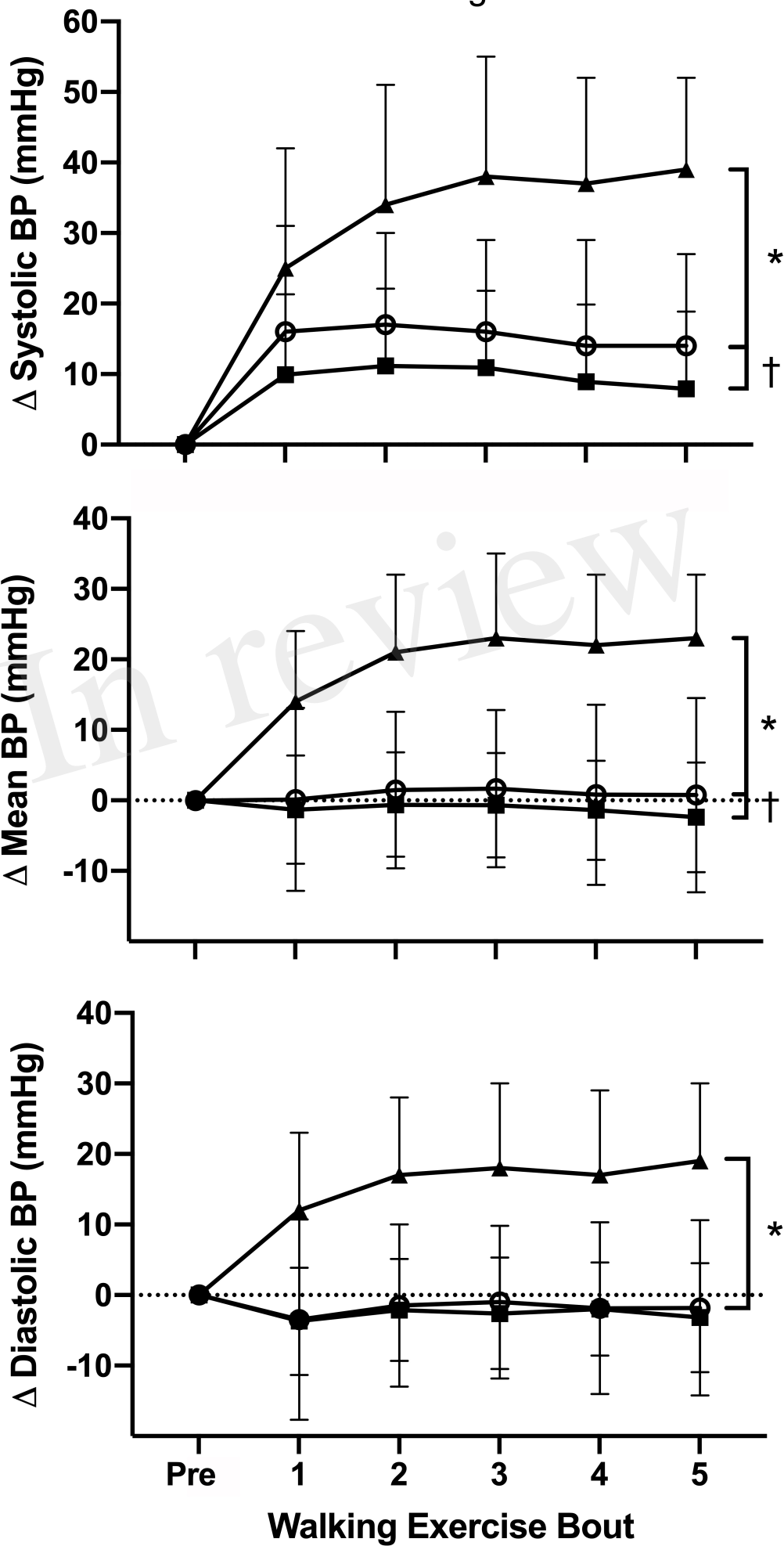


Figure 2.JPEG

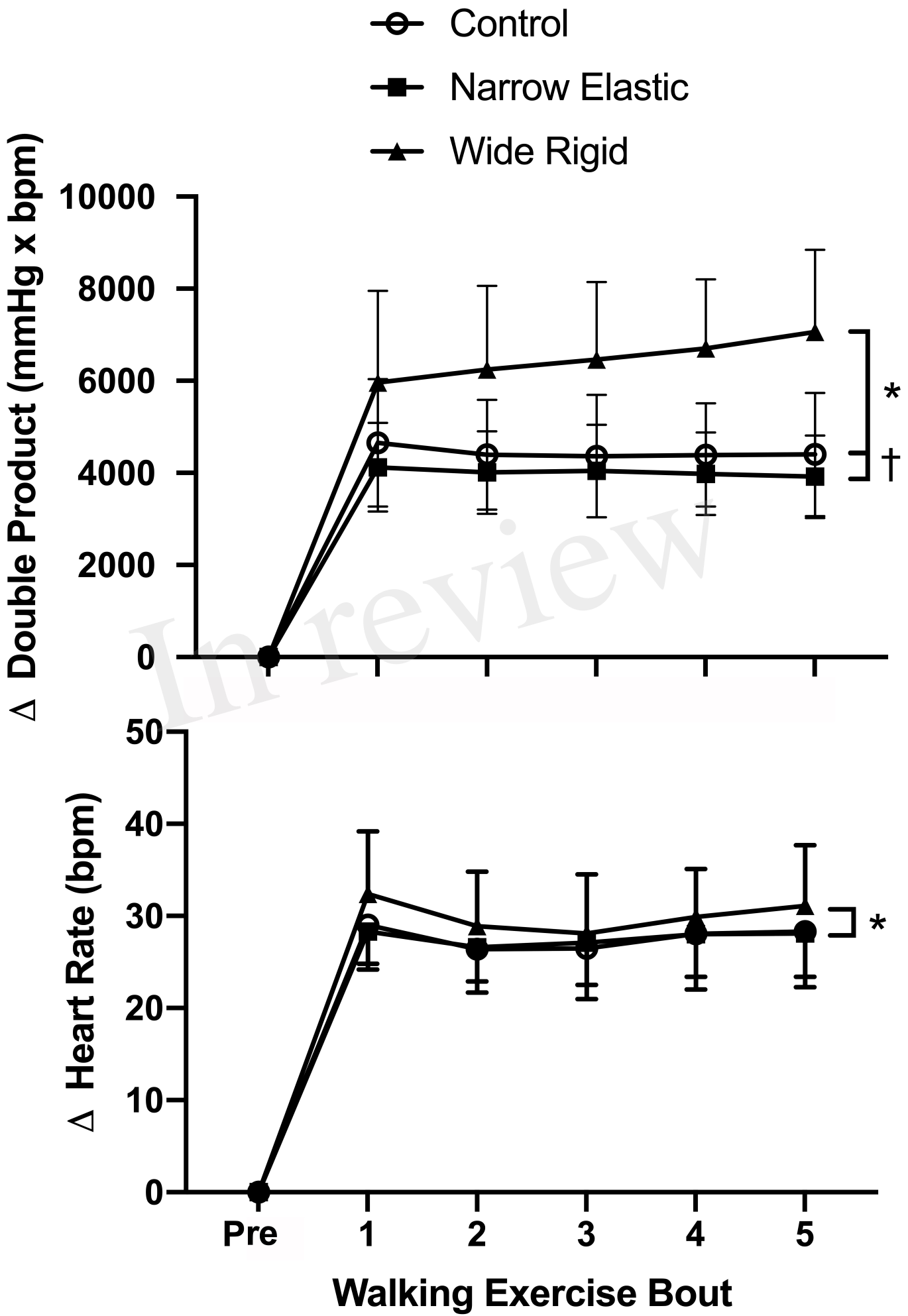


Figure 3.JPEG

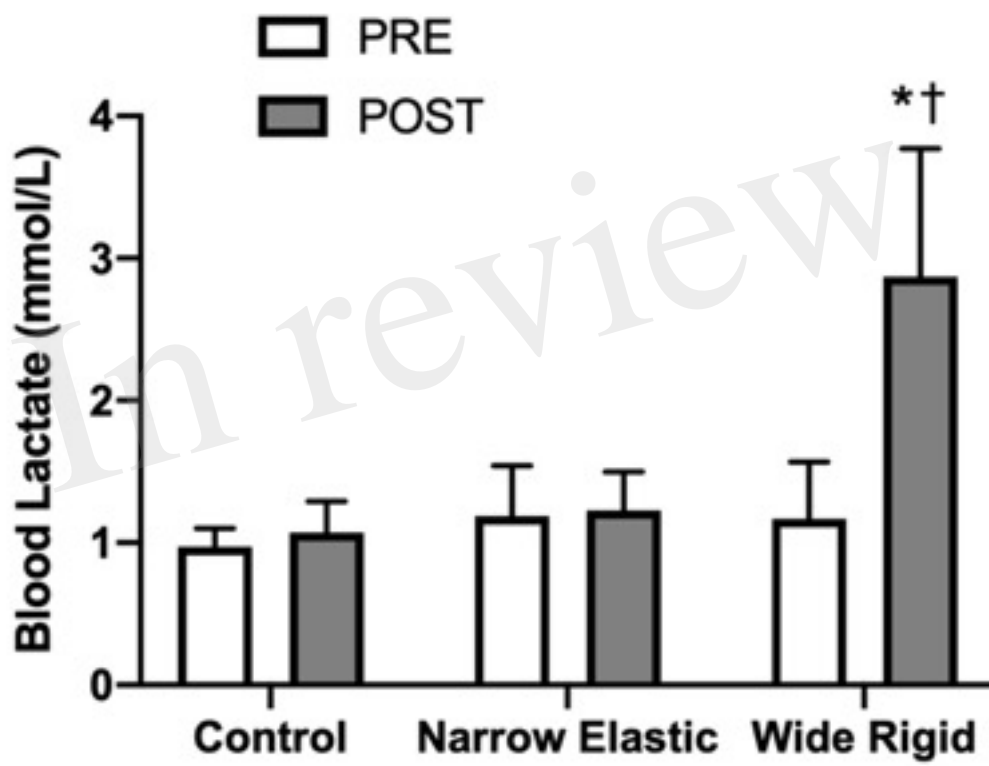


Figure 4.JPEG

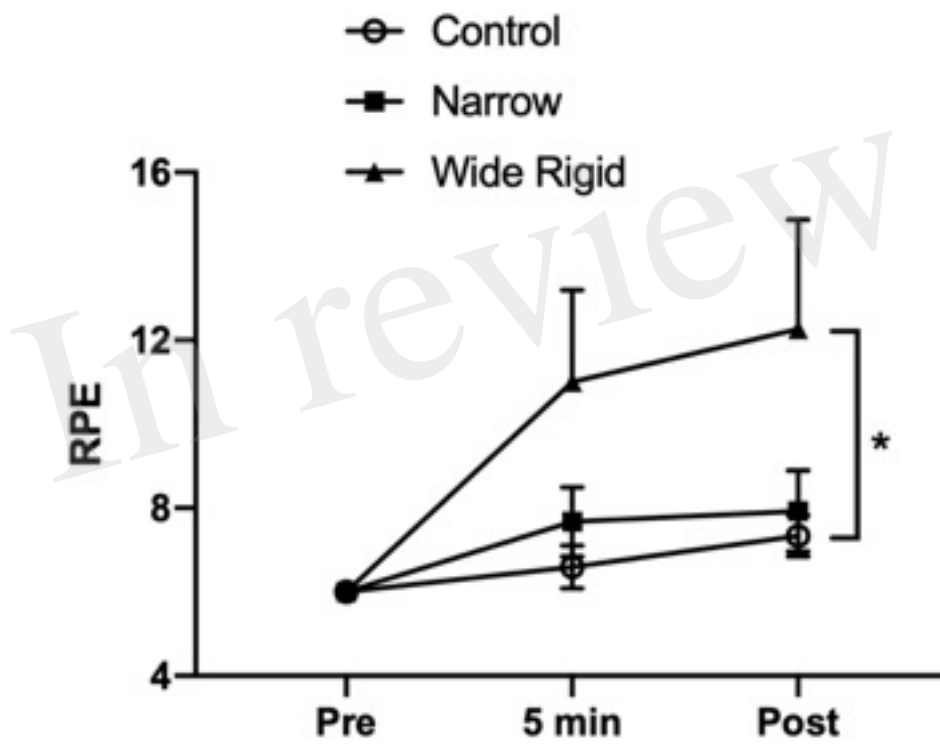


Figure 5.JPEG

