- 1 Title: Why BFR cuff features are an important methodological consideration- A short
- 2 commentary on "Cerebral cortex activation and functional connectivity during low-load
- 3 resistance training with blood flow restriction: An fNIRS study"
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- 34 This is a preprint and has not been peer-reviewed.
- 3536 Please cite:
- 37 Rolnick, N., Clarkson, M., Hughes, L., Korakakis, V., De Queiros, V., Patterson, S. D., Buckner,
- 38 S., Werner, T., Da Cunha Nascimento, D., Stray-Gundersen, S., Kamis, O., Thoelen, M.,
- 39 Kimbrell, K., & Jacobs, E. (2024). Why BFR cuff features are an important methodological
- 40 consideration: A short commentary on "Cerebral cortex activation and functional connectivity
- 41 during low-load resistance training with blood flow restriction: An fNIRS study." Sportrxiv.
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48	Before reading the article, we want to reiterate our desire to have an open and honest
49	dialogue with Jia et al. (2024) regarding the content of their manuscript. We wrote this
50	letter to the editor and Dr. Jeremy Loenneke, PhD – the editor for this publication - felt
51	that this commentary was worthy of publication to begin the discussion surrounding
52	the article. However, he was denied of proceeding forward to publication by the
53	PLOSOne policy of not publishing letters to the editor/commentaries. Moreover, the
54	only place where we could leave a commentary on the article is in the "Public
55	comments" section. Unfortunately, it is not working due to undisclosed
56	circumstances and there is no timeline for when it will be back online. Nonetheless,
57	we are extremely disappointed in PLOSOne's policy to not allow for open and honest
58	academic discourse on their publications. We hope that in the future PLOSOne allows
59	for more open discussions on the articles they publish.
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94 We read with great interest the recent study titled "Cerebral cortex activation and functional

- 95 connectivity during low-load resistance training with blood flow restriction: An fNIRS study"
- 96 (Jia et al., 2024). The study adds to our limited understanding of the cerebral demands of blood
- 97 flow restriction (BFR) exercise and the potential role of applied pressure. The authors examined
   98 cerebral oxygenation levels following squat exercise performed at 30% of one repetition
- maximum, with bilateral BFR applied at 150, 250, and 350 mmHg using the B-Strong cuffs (B-
- 100 Strong, USA). The authors noted enhanced cerebral oxygenation levels in many cortical regions
- 101 which dropped sharply when 350 mmHg was applied. In addition, they also found the existence
- 102 of an interaction effect of pressure on cortical activation in the primary motor cortex, pre-motor
- 103 cortex, and supplementary motor cortex whereas there was a less pronounced effect in the
- 104 dorsolateral prefrontal cortex. The authors should be commended for their pioneering
- 105 investigation into the relationship between applied BFR pressures and cortical demands.
- 106 However, we wish to bring up some methodological concerns and considerations regarding the
- 107 cuff utilized as well as the way that pressure was applied in data collection and speculate on its
- 108 potential impact and influence on the ultimate outcomes as calculated and reported in this study. 109
- 110 In the last decade, BFR has grown in popularity in multiple practice settings (Scott et al., 2023).
- As a result of this popularity, BFR cuff manufacturers have begun to produce different types of
  BFR equipment and incorporate device features that can impact the acute and/or longitudinal
  responses to BFR exercise (Rolnick et al., 2023). Features such as autoregulation of applied BFR
- 114 pressures during exercise (L. Hughes et al., 2023). Teatures such as autoregulation of applied BF
- 115 (Buckner et al., 2017; Loenneke et al., 2012) or changes in the bladder design that houses the air
- 116 that is applied to the limb (Dancy et al., 2012) by changes in the bladder design that is
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118 Jia et al. (2024) utilized the B-Strong cuff, a multi-chambered BFR cuff that is designed to avoid significant arterial occlusion to promote user safety during its application (Rolnick & Cerqueira, 119 120 2021). These are distinct from single air bladder (e.g., a traditional tourniquet) cuffs that are 121 designed to determine a personalized pressure (Limb occlusion pressure, LOP) during BFR 122 exercise (Patterson et al., 2019). LOP has been defined as the minimum applied pressure needed 123 to fully occlude arterial and venous blood flow to an extremity, and provides a way to standardize BFR application (Patterson et al., 2019). Personalizing the pressure application has 124 125 been recommended in clinical practice and research because it allows for similar comparisons 126 between participants and can assist practitioners in implementing applied pressures that influence 127 relevant physiological outcomes. LOP values are largely predicated on the BFR cuff width and 128 each participant's resting blood pressure, limb circumference, and body position (Graham et al., 129 1993; Luke Hughes et al., 2018; Loenneke et al., 2013; Sieljacks et al., 2018). Relativizing the

- applied pressure for each individual using the LOP approach ensures these participantcharacteristics are taken into consideration and can provide a better estimation of the applied
- 132 pressure and the extrapolation and comparison of findings between conditions and laboratories.
- 133 While the absolute amount of pressure applied to each participant may vary significantly when
- 134 standardizing the pressure application to a percentage of LOP between cuffs of different sizes,
- the physiological stimulus appears similar (Loenneke et al., 2012). For example, 250 mmHg
- applied pressure to one individual may be complete occlusion whereas it may only be partial
- 137 occlusion to another individual based upon individual characteristics. Therefore, an important

- physiological responses, including cerebral oxygenation, is utilizing cuffs and methods that canrelativize the applied BFR pressure.
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142 As the primary goal of the current study was to determine the pressure-dependent relationship to 143 cortical activation and cerebral oxygenation, the use of a multi-chambered cuff without a 144 standardized method to relativize the applied pressure could impact any potential effect observed 145 from increasing pressure compared to a single-chambered bladder BFR cuff. Prior research has 146 shown arterial blood flow only begins to be modified from resting conditions with greater than 147 350 mmHg of applied pressure when using multi-chambered BFR cuffs (Citherlet et al., 2022). 148 Conversely, pressures as low as ~86 mmHg (40% LOP in this particular study) were shown to 149 modulate blood flow from resting conditions in the Hokanson device (Citherlet et al., 2022). It is 150 tempting to suggest that 350 mmHg with a multi-chambered bladder BFR cuff and 40% LOP 151 with a single-chambered bladder BFR cuff provide a similar physiologic stimulus. However, 152 without instituting methods to relativize the applied pressures in the multi-chambered cuff, it is 153 difficult to know. Nonetheless, if this comparison is true, it would suggest that a pressure of 350 mmHg in a multi-chambered BFR cuff, which is on the low end of the recommendations for 154 applied pressure during BFR exercise using single-chambered cuffs (40-80% LOP) (Patterson et 155 156 al., 2019), alters cortical activation and cerebral oxygenation. Given the standard application of a 157 fixed pressure in lieu of a relativized application, participants in Jia et al. (2024) were likely 158 exercising at different levels of pressure relative to their LOP, creating uncertainty around the 159 findings and its translation to practice.

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161 We recommend that future studies either consider personalizing to a %LOP or standardize the 162 cuff fitting pressure when using multi-chambered cuffs and attempting to elucidate pressuredependent changes in outcome measures. At the very least, individual features, such as the 163 participant's resting blood pressure and limb circumference should be reported to provide greater 164 165 context. Some research shows that multi-chambered cuffs can be personalized (Machek et al., 166 2022), so use of this cuff design feature in research studies to explore the role of applied pressure 167 is not necessarily a methodological flaw but does require additional steps to contextualize (e.g., measurement of LOP). Conversely, applying an arbitrary amount of pressure for each condition 168 169 reduces generalizability and limits the strength of the findings. This is particularly important 170 considering the only pressure condition capable of reducing cerebral oxygenation and activity in 171 the Jia et al. (2024) study represents not only the minimum pressure threshold needed to decrease 172 resting arterial blood flow with the B-Strong cuffs (Citherlet et al., 2022), but also was the maximum pressure examined. Further, the authors did not mention this arbitrary pressure 173 application approach as a limitation. The authors mentioned the method of compression and the 174 175 material of the compression band, but we assert that the multi-chambered bladder design is the biggest limitation to this line of BFR research on pressure-dependent relationships. In summary, 176 177 we commend the authors for investigating a novel component of BFR training but hope that highlighting the potential impact of cuff design and BFR methodology can have on the BFR 178 179 stimulus spurs careful consideration of these factors and generates more standardization of BFR 180 application in research settings. 181

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## Author Contributions

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187 Nicholas Rolnick wrote the first draft. The remaining authors provided critical commentary and
188 feedback on subsequent drafts and all authors approved its final form.

- Competing interests:

Nicholas Rolnick is the founder of THE BFR PROS, a BFR education company that provides BFR training workshops to fitness and rehabilitation professionals across the world using a variety of BFR devices. Kyle Kimbrell is a clinical instructor for Owens Recovery Science, a BFR education company that distributes the Delfi Personalized Tourniquet Device. Sten Stray-Gundersen is a clinical instructor for B-Strong Training Systems. Vasileios Korakakis, Luke Hughes, Dahan Da Cunha Nascimento, Mathias Thoelen, and Ewoud Jacobs are clinical instructors for blood flow restriction and deliver education courses to practitioners with no company affiliation. The other authors declare no potential or actual conflicts of interest.

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