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Orienteering combines vigorous-intensity exercise with navigation to
improve human cognition and increase brain-derived neurotrophic
factor

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25 **Abstract**

26 Exercise enhances aspects of human cognition, but its intensity may matter. Recent animal
27 research suggests that vigorous exercise, which releases greater amounts of lactate, activates
28 more brain-derived neurotrophic factor (BDNF) in the hippocampus and, thus, may be optimal
29 for supporting cognitive function. The cognitive benefits of exercise may be further augmented
30 when combined with cognitive training. The sport of orienteering simultaneously combines
31 exercise with spatial navigation and, therefore, may result in greater cognitive benefits than
32 exercising only, especially at vigorous intensities. The present study aimed to examine the
33 effects of an acute bout of orienteering at different intensities on cognition and BDNF compared
34 to exercising only. We hypothesized that vigorous-intensity orienteering would increase lactate
35 and BDNF and improve cognition more than moderate-intensity orienteering or vigorous
36 exercise alone. Sixty-three recreationally active, healthy young adults ($M_{\text{age}} = 21.10 \pm 2.75$ years)
37 with no orienteering experience completed a 1.3 km intervention course by navigating and
38 exercising at a vigorous (80-85% of heart rate reserve) or moderate (40-50% of heart rate
39 reserve) intensity or exercising vigorously without navigation. Exercise intensity was monitored
40 using peak lactate, heart rate and rating of perceived exertion. Serum BDNF was extracted
41 immediately before and after the intervention. Memory was assessed using the Mnemonic
42 Similarity Task (high-interference memory) and the Groton Maze Learning Test (spatial
43 memory). Both exercising and orienteering at a vigorous intensity elicited greater peak lactate
44 and increases in BDNF than moderate-intensity orienteering, and individuals with higher peak
45 lactate also had greater increases in BDNF. High-interference memory improved after both
46 vigorous-intensity interventions but did not improve after the moderate-intensity intervention.
47 Spatial memory only increased after vigorous-intensity orienteering, suggesting that orienteering

48 at a vigorous intensity may particularly benefit spatial cognition. Overall, the results demonstrate
49 the benefits of vigorous exercise on human cognition and BDNF.

50

51 **Introduction**

52 As the brain ages, atrophy often outpaces plasticity, resulting in neurodegeneration and
53 cognitive decline. Some brain regions are more susceptible to age-related decline than others,
54 and the hippocampus is one of them [1]. After the age of 55 years old, the hippocampus
55 atrophies at a rate of about 0.5% percent per year but progresses at twice that rate after the age of
56 70 years old [2] and nearly eight times that rate for individuals with Alzheimer's disease [3].
57 This selective and severe hippocampal degeneration can impair critical hippocampal functions
58 such as learning, memory, and spatial cognition and may compromise independent living [4].
59 Age is the greatest risk factor for dementia, and as the world's population ages, dementia rates
60 are predicted to climb sharply to affect over 152 million people by 2050 [5]. With no known cure
61 for dementia, preventative measures that can help to stave off age-related cognitive decline are
62 essential.

63 Exercise is one way to boost plasticity; however, emerging evidence suggests that not all
64 forms of exercise are as effective. Vigorous exercise tends to evoke greater increases in plasticity
65 through its stimulation of brain-derived neurotrophic factor (BDNF), a neurotrophic factor that
66 supports the growth, function and survival of brain cells [6,7]. Vigorous exercise has been
67 associated with memory improvements in both younger [8] and older adults [9]. New research
68 from animal models suggests that muscle-to-brain signalling during vigorous exercise is
69 mediated by l-lactate (herein referred to as lactate), a product of pyruvate metabolism under
70 anaerobic conditions that accumulates with increasing exercise intensity [10] and increases

71 exponentially beyond the lactate threshold of ~ 4mmol/L of lactate in untrained adults [11,12].
72 Although lactate has historically and erroneously been considered an inert metabolic waste [13],
73 recent evidence points to its importance as both a fuel source [14] and an activator of BDNF
74 [15–18] with rapid effects. Mere minutes after the initiation of vigorous exercise, lactate-
75 activated BDNF has the potential to facilitate long-term potentiation within existing neural
76 synapses to enhance neuroplasticity [19]. In this way, lactate accumulation during an acute bout
77 of vigorous exercise may explain why acute exercise can immediately enhance certain cognitive
78 functions [20]. To date, most research on the lactate-cognition connection has been done in
79 animal models; only a few studies demonstrated the association in humans [10,21–23].
80 Therefore, a primary objective of the present study was to examine the role of lactate in muscle-
81 to-brain signalling on BDNF and cognition in humans.

82 We also wanted to examine whether the effects of vigorous exercise could be enhanced
83 when simultaneously combined with a cognitively challenging task. During the process of
84 neurogenesis, exercise predominantly impacts the proliferation of newborn neurons in the
85 dentate gyrus, whereas cognitive training predominantly impacts the maturation and survival of
86 those newborn brain cells [24]. Consequently, when combined, there is the potential for additive
87 effects. Indeed, simultaneous exercise-cognition interventions in older adults improves cognition
88 more than sequential interventions or cognitive training alone [25]. For example, older adults
89 who engaged in spatial navigation while treadmill walking experienced enhancements in their
90 spatial cognition more than older adults who only walked on the treadmill. Moreover, after four
91 months of training, walkers saw a decrease in hippocampal volume, whereas navigators
92 maintained a consistent volume, suggesting that there are added neurogenic benefits of
93 combining exercise with navigation [26]. While intriguing, the mechanisms underlying these

94 augmentative effects in humans are unclear, especially concerning the role that lactate and
95 BDNF may play in promoting cognition, and testing those associations was the primary aim of
96 this study.

97 For our simultaneous exercise-cognition training, we used the sport of orienteering,
98 which naturally and simultaneously integrates exercise with spatial navigation and, therefore,
99 may be an optimal way to combine exercise and cognitive training to target hippocampal
100 plasticity and function [27]. The sport of orienteering requires the athlete to navigate through a
101 series of checkpoints across an unknown terrain as fast as possible using only a topographical
102 map and a compass [28]. Through focused attention and quick deduction of key information,
103 highly skilled orienteers use spatial information and mental representations of an environment to
104 navigate efficiently through space [28,29], which is a critical function of the hippocampus [30].
105 Atrophy of the hippocampus impairs spatial navigation [31], and in cases of advanced AD,
106 severe hippocampal degeneration renders the hippocampus unable to create, store, or use mental
107 maps for wayfinding [32], causing disorientation even in familiar environments, a condition
108 known as topographical disorientation [33,34]. In line with the “use it or lose it” hypothesis [35],
109 modern-day dependencies on vehicles for transport and passive navigation guided by Global
110 Positioning Systems (GPS) cause most humans to underutilize their wayfinding abilities, leading
111 to spatial memory deficits [36] and a reduced sense of direction [37] which orienteering has the
112 potential to rescue. Moreover, to navigate through their environment, orienteers engage in
113 various sensorimotor processes, and therefore, concepts of embodied cognition may also be
114 relevant [38].

115 Indeed, our prior research revealed that orienteering experts aged 18-87 reported superior
116 navigational strategies and better spatial memory than non-orienteering controls [27]. This recent

117 observation resembles earlier research on London taxi drivers who, compared to controls, had a
118 higher degree of navigational competency [39]. The taxi drivers also had a larger posterior
119 hippocampus, a brain region primarily involved in supporting better visuospatial cognition,
120 whose larger size was associated with greater years of experience [39–41]. However, not all parts
121 of their hippocampus were larger; the anterior hippocampus, historically understood for its role
122 in mediating episodic memory, was smaller in taxi drivers compared to controls, suggesting a
123 trade-off between spatial and episodic memory that may be dependent on the training experience.
124 Notably, the same trade-off was not seen with orienteering in that expert orienteers reported
125 better spatial memory but not worse episodic memory to controls [27]. The simultaneous
126 integration of exercise with navigation may be preventing the trade-off [27]. To date, only a
127 handful of studies have examined the effect of orienteering training on cognition [42–44]; most
128 have examined spatial cognition, and none have manipulated its intensity or examined lactate
129 and BDNF.

130 Therefore, the present study aimed to examine the effects of orienteering at different
131 exercise intensities (vigorous versus moderate) compared to vigorous intermittent exercise only
132 on lactate, BDNF and different aspects of hippocampal-dependent memory. We hypothesized
133 that the vigorous-intensity interventions would increase lactate more than the moderate-intensity
134 intervention, resulting in a greater increase in BDNF and memory. Given the potential for
135 additive effects of exercise-cognition training, we hypothesized that orienteering at a vigorous
136 exercise intensity would elicit larger gains in BDNF and memory compared to orienteering at a
137 moderate intensity or vigorous exercise alone.

138

139 **Methods**

140 **Participants**

141 Sixty-three participants (n = 41 female) who were healthy young adults ($M_{\text{age}} = 21.10$,
142 $SD = 2.75$, range = 18-30) were recruited to the study using self-referral based on the criteria of
143 being aged 18-30 years old and recreationally active (i.e., achieving at least 150 minutes up to
144 4.5 hours of recreational moderate-to-vigorous physical activity per week, as confirmed using the
145 Physical Activity and Sedentary Behaviour Questionnaire; [45]. Recruitment was ongoing
146 between July 2022 to May 2023. Participants were only included if they had engaged in
147 orienteering from zero to a maximum of five times, a criterion based on previous research where
148 an “orienteer” was defined as someone with at least six sessions of orienteering training [46].
149 Participants were screened to ensure eligibility using the following self-reported criteria: 1) no
150 diagnosis of a neurological disorder or major health condition, 2) English language fluency, and
151 3) no colour blindness. Written informed consent was obtained through an online questionnaire.
152 Participants were randomized into one of three groups: 1) moderate-intensity orienteering (n =
153 22), 2) vigorous-intensity orienteering (n = 21) and 3) vigorous-intensity exercise (n = 20), as
154 described below. Participants received an honorarium of thirty Canadian dollars for their time.
155 This study was reviewed and approved by the Hamilton Integrated Research Ethics Board
156 (#14560) before recruitment and data collection.

157 **Materials and procedure**

158 **Baseline questionnaires**

159 Following randomization and before the in-lab session, participants completed an online
160 questionnaire (LimeSurvey software) to collect demographic information (see S1 Appendix).

161 All participants then completed the Physical Activity and Sedentary Behaviour
162 Questionnaire [45] to assess their average weekly amount of moderate-to-vigorous aerobic

163 exercise. The total activity amount was determined by multiplying the average length of an
164 exercise session by the average number of active days (minutes/week).

165 The Navigational Strategy Questionnaire (NSQ) was used to assess participants' baseline
166 navigational tendencies [47]. Using a 5-point Likert scale, participants rated 44 items
167 corresponding to three different navigational strategies: allocentric spatial processing, egocentric
168 spatial processing, and procedural processing. For each strategy, an average score was
169 calculated.

170 Baseline autobiographical memory was assessed using the Survey of Autobiographical
171 Memory (SAM; [48]). In the SAM, subjective memory is evaluated across 26 items which are
172 answered using a 5-point Likert scale. Each item is weighted and summed to obtain an average
173 for four memory domains including episodic, spatial, semantic, and future memory. In this study,
174 we examined episodic and spatial memory specifically.

175 **Lab-based baseline measurements**

176 In the lab, before the intervention, the participant's height (centimetres), weight
177 (kilograms), and waist circumference (in centimetres and taken from the anterior-superior iliac
178 spine upon exhalation) were measured by a trained researcher.

179 Resting heart rate (HR_{Rest}) was determined using a wetted Polar HR-10 chest heart rate
180 (HR) monitor synchronized to a Polar Pacer Pro watch (Polar Electro Canada, Lachine, Quebec).
181 The lowest HR value recorded in the final two minutes of a 12-minute supine resting period was
182 used.

183 Maximum heart rate (HR_{Max}) was estimated using the equation $HR_{Max} = 208 - (0.7 * age)$
184 [49]. HR_{Rest} and HR_{Max} were used to calculate their exercising heart rate zones for the
185 intervention using the percent of heart rate reserve (HRR) and the equation $(HR_{Max} - HR_{Rest}) *$

186 (intensity) + HR_{Rest} . For the moderate-intensity orienteering group, the exercise intensity range
187 was calculated as 40-50% of HRR, and 80-85% of HRR was used for the vigorous-intensity
188 orienteering and vigorous-intensity exercise groups.

189 Estimates of VO_2 peak were calculated using the *WorldFitnessLevel.org* website [50].
190 Participants were asked to respond to the website questions as accurately as possible and input
191 their anthropometric and HR measurements.

192 **Intervention measures of exercise intensity**

193 During the intervention, HR, ratings of perceived exertion (RPE) and lactate were
194 recorded at the middle and end of the intervention course and 10 minutes post-intervention. The
195 highest of these values was analyzed. Heart rate was recorded using the Polar HR-10 monitor.
196 RPE was captured using the Borg 1-20 Scale [51]. Lactate was measured from a sample of whole
197 blood obtained from the fingertip using the Lactate Plus portable analyzer (Nova Biomedical,
198 Waltham, MA).

199 **Pre- and post-intervention measurements**

200 Before the intervention, the cognitive testing was completed before obtaining a serum
201 sample for BDNF. Following the intervention, the blood sample was collected within 10 minutes
202 of finishing the intervention course and was followed by cognitive testing.

203 For BDNF, three-hour fasted samples of venous blood were obtained from a vein in the
204 antecubital fossa. Samples were collected into BD Vacutainer SST tubes (BD, Franklin Lakes,
205 NJ), chilled on ice, allowed to clot for a minimum of 45 minutes following sample collection and
206 then centrifuged at $1000 \times g$ for 15 minutes at $4^\circ C$. For all samples, $300\mu L$ of supernatant was
207 collected to obtain serum, aliquoted into microtubes, and stored immediately at $-20^\circ C$ until
208 analysis. The concentration of serum BDNF was quantified using a sandwich Biosensis Mature

209 BDNF Rapid™ ELISA Kit (Biosensis Pty Ltd, Thebarton, Australia). Samples were diluted
210 100x and were run in duplicate. Using a BioTek SynergyMx spectrophotometer, absorbance was
211 measured at 450 nm and analyzed using Gen 5 1.11 Software (BioTek Instruments Inc.,
212 Winooski, VT). Select samples whose concentration fell above the standard curve of the
213 preliminary analysis were re-analyzed using a 125x dilution and the same protocol.

214 Memory was tested in two ways. First, memory was tested using Kirwan and Stark's
215 Mnemonic Similarity Task [52–54], a modified object recognition task that places a large
216 emphasis on high-interference memory and hippocampal function. The Mnemonic Similarity
217 Task begins with a study phase in which participants are shown a series of images of 60
218 everyday objects displayed on the screen for two seconds and must classify whether the image is
219 an 'indoor' or an 'outdoor' item. This is immediately followed by a test phase, in which
220 participants are shown 20 'repeat' images (correct response = "old"), 20 'lure' images that are
221 highly similar but not identical to a previous image (correct response = "similar"), and 20
222 completely new, 'foil,' images (correct response = "new") and asked to classify them. The
223 Mnemonic Similarity Task has two measures which provide a valuable distinction between
224 hippocampal-dependent high-interference memory and recognition memory. The "lure
225 discrimination index" is a measure of high-interference memory, calculated as $[p(\text{"Similar"} |$
226 $\text{Lure image}) - p(\text{"Similar"} | \text{Foil image})] \times 100$, and reflects one's ability to correctly classify
227 'lure' items as "similar". High-interference memory relies on the ability to remember specific
228 details during encoding [55], which is dependent on the function of the hippocampus and is
229 associated with hippocampal neurogenesis [54]. The second measure of the Mnemonic Similarity
230 Task is general "recognition memory", defined as the ability to correctly label a 'repeat' image
231 as "old," $[p(\text{"Old"} | \text{Repeat image}) - p(\text{"Old"} | \text{Foil image})] \times 100$. Recognition memory does

232 not require participants to distinguish between highly interfering memories. It is less impacted by
233 exercise and, therefore, theorized to be less dependent on hippocampal neurogenesis [56,57]. The
234 Mnemonic Similarity Task was administered before and after the intervention with different
235 stimulus sets, and the order of each set was counterbalanced.

236 The second test of memory assessed spatial learning and memory using a computerized
237 version of the Groton Maze Learning Test, adapted from the Milner Maze [58]. The 2D maze
238 consists of a 28-step pathway hidden beneath a 10x10 grid of grey tiles that is revealed by
239 clicking on adjacent tiles within the matrix using a mouse. If a correct tile in the sequence is
240 selected, the tile briefly turns green, a rewarding auditory tone is played, and the participant can
241 select a new tile. If an incorrect tile is selected, the tile briefly turns red, an incorrect auditory
242 signal is played, and the participant must click on the previously correct tile before choosing a
243 new tile. In the learning phase, participants complete the same maze five times in a row as fast as
244 possible. To test delayed memory, participants complete the same maze once more following a
245 10-minute break. A Maze Efficiency Index [59] was calculated using the equation $\text{Maze Efficiency Index} = \text{number of correct moves per second} / \text{Log}_{10}(\text{time of the trial})$. Mean Maze
246 Efficiency Index was calculated for the learning phase by averaging across the five learning trials
247 whereas the Maze Efficiency Index for the test phase consisted of performance on the single test
248 trial. The Groton Maze Learning Test was administered before and after the intervention with
249 randomized maze sequences using computer software.

251 **The intervention**

252 The intervention started with a practice phase during which all participants completed a
253 500-meter outdoor practice course on the McMaster campus that included six orienteering
254 checkpoints to get warmed up and familiarized with the intervention procedures. Participants

255 learned how to read their HR on the Polar Pacer Pro GPS watch and maintain their pace so that
256 their HR remained in the target range. Participants in the orienteering groups (both moderate and
257 vigorous intensity) were taught how to use the orienteering map legend, orient their map, use the
258 map to plan a route and locate checkpoints, and re-locate themselves should they make an error.
259 Then, to simulate the intervention, the orienteering groups located the first three checkpoints at a
260 light walking pace of 30-40% of HRR with the help of a researcher. For the fourth checkpoint,
261 participants were encouraged to locate the checkpoint independently. For the final two
262 checkpoints, participants located the checkpoints on their own and at their target intensity
263 (moderate-intensity: 40-50% of HRR) or running (vigorous-intensity: 80-85% of HRR) pace. In
264 contrast, participants in the vigorous exercise only group did not actively navigate. Instead, they
265 followed a researcher around the 500 m course, beginning at a walking pace of 30-40% of HRR
266 for the first four checkpoints and at their target intensity of 80-85% of HRR for the final two
267 checkpoints.

268 Immediately after the practice phase, all participants were led to the start location of the
269 intervention course by a researcher, and the Polar Pacer Pro GPS watch was started to track the
270 participant's HR and route. All participants completed the intervention course, which was
271 approximately 1.3 kilometers and consisted of 10 checkpoints around the McMaster University
272 campus according to their intervention condition. Those in the orienteering groups navigated to
273 the checkpoints using the map at either a moderate (40-50% of HRR) or vigorous intensity (80-
274 85% of HRR) along any route they chose. For safety reasons, a researcher silently followed
275 participants during the intervention. For participants who were severely lost or had ventured
276 outside the bounds of the orienteering map, the researcher informed them of their current
277 location to ensure their safety but did not provide any additional information that would alter

278 their navigational decisions. In contrast, those in the exercise only group exercised at a vigorous
279 intensity (80-85% of HRR) but did not engage in orienteering. Instead, a member of the research
280 team led the participant along the most efficient route.

281 All participants were responsible for tracking their HR at each checkpoint and were
282 instructed to adapt the pace or pause until their HR returned to the target zone for a maximum of
283 one minute. At the midpoint and finish checkpoints, a second researcher recorded HR, RPE and
284 blood lactate.

285 **Statistical analysis**

286 All data were analyzed using SPSS (IBM SPSS Statistics for Macintosh, version 28.0;
287 IBM Corp., Armonk, NY). For all study variables, descriptive statistics were computed.
288 Normality was assessed using skewness, kurtosis, and visual inspection of histograms. Data were
289 screened for outliers using visual inspection of boxplots. For BDNF, cases were removed if
290 BDNF concentration was above the standard curve, in which seven cases were removed
291 (moderate orienteering = 1, vigorous orienteering = 4, vigorous exercise = 2). For the Mnemonic
292 Similarity Task, three cases were removed as the difference in the percent corrected and raw
293 score for appropriate key use was >8% (moderate orienteering = 2, vigorous orienteering = 1).
294 Cases were also removed due to programming errors with the cognitive test software (Mnemonic
295 Similarity Task: moderate orienteering = 1, vigorous orienteering = 1; Groton Maze Learning
296 Test: moderate orienteering = 1, vigorous orienteering = 1) and because of errors in GPS data
297 recording (vigorous orienteering = 2). Only complete cases were analyzed for each variable. All
298 tests were computed with an alpha criterion of .05 and a 95% confidence interval.

299 **Potential covariates and manipulation checks**

300 To test for potential covariates, a one-way analysis of variance (ANOVA) was used to
301 assess group differences in all demographic variables, weekly physical activity, spatial
302 navigation tendencies and autobiographical memory, as well as pre-intervention differences in
303 BDNF and cognition. To ensure that our intervention was adequate in reaching the desired
304 exercise intensity, one-way ANOVA tests were computed for peak HR, peak RPE and peak
305 blood lactate between groups. For blood lactate, a Kruskal-Wallis Means Ranks Test was used to
306 confirm that the proportions of those above or below the lactate threshold of 4mmol/L differed
307 by group, thus indicating that our intervention was adequate in reaching the desired exercise
308 intensity.

309 **Primary outcome measures**

310 All primary outcome variables (BDNF, high-interference memory (lure discrimination
311 index), recognition memory and spatial learning and memory efficiency) were analyzed using
312 separate 2 x 3 mixed model ANOVAs with a within-subjects factor of time (pre, post) and
313 between-subjects factor of group (moderate orienteering, vigorous orienteering, vigorous
314 exercise). *A priori* one-sample *t*-tests (one-tailed) were computed to evaluate the pre- to post-
315 intervention increases in BDNF and memory for each group with Hedge's correction. Post hoc
316 analyses of any between-group comparisons were performed with Bonferroni correction.
317 Spearman's correlation was used to evaluate the relationship between peak lactate and percent
318 change in BDNF.

319 To further explore the relationship between peak lactate, percent change in BDNF and
320 cognitive function, we computed a composite cognitive score was calculated by averaging the z-
321 scores for the post-minus-pre change score values for each of our cognitive measures (high
322 interference memory, recognition memory, Groton Maze learning efficiency, and Groton Maze

323 test efficiency). Then, we performed an exploratory analysis using Spearman's correlation to
324 evaluate the relationship between the composite cognition score with peak lactate and percent
325 change in BDNF. Finally, we conducted a partial Spearman's correlation to determine whether
326 the association between composite cognition score and peak lactate was diminished after
327 controlling for the percent change in BDNF.

328 **Secondary outcome measures**

329 An exploratory analysis was done to quantify differences in the navigational performance
330 of the two orienteering groups. The distance travelled by each of the orienteering groups
331 (moderate orienteering, vigorous orienteering) was compared to the vigorous exercise group
332 which, by design, travelled the most efficient route. A 2 x 3 mixed model ANOVA with a
333 within-subjects factor of course half (start to midpoint, midpoint to finish) and between-subjects
334 factor of group (moderate orienteering, vigorous orienteering, vigorous exercise) was used to
335 identify group differences in distance travelled as indicated by the Polar Pacer Pro GPS watch.
336 Post hoc analyses used a Bonferroni correction. Spearman's correlation was used to determine
337 existing associations between the total distance travelled and subjective measures of spatial
338 processing/navigation and memory and for baseline measures of cognitive function for the two
339 orienteering groups.

340

341 **Results**

342 **Participants**

343 Table 1 reports descriptive statistics of key baseline variables across groups. Ninety-two
344 percent (n = 58/63) of participants were students at McMaster University. Participants did not
345 differ in pre-exercise measures of high-interference or recognition memory or in spatial learning

346 and memory. However, pre-intervention BDNF levels were higher for the moderate orienteering
 347 group than the vigorous orienteering or vigorous exercise groups ($p < .001$) (Table 2). Univariate
 348 ANOVA tests confirmed no other baseline differences between groups (Table 1).

349

350 **Table 1. Descriptive Statistics Between Intervention Groups**

	Moderate Orienteering	Vigorous Orienteering	Vigorous Exercise
<i>n</i>	22	21	20
Age (years)	20.48 ± 2.34	21.76 ± 3.36	21.05 ± 2.46
Age Range (years)	18 - 28	18 - 30	18 - 26
Sex (F/M)	14/8	13/8	14/6
Height (cm)	170.16 ± 7.52	166.79 ± 8.53	169.40 ± 9.30
Weight (kg)	67.38 ± 10.53	68.91 ± 12.64	63.91 ± 14.89
WC (cm)	82.52 ± 7.15	83.80 ± 7.72	81.97 ± 10.17
Aerobic Physical Activity (min/week)	172.74 ± 91.12	172.62 ± 99.64	198.25 ± 103.76
Predicted VO₂Max (mL/kg/min)	49.23 ± 6.58	50.19 ± 6.23	48.95 ± 5.51
Education			
< Secondary	0%	0%	5%
Secondary	82%	57%	75%
Post-Secondary	18%	24%	5%
Post-Graduate	0%	19%	15%
McMaster Student (No/Yes)	1/21	2/19	2/18
McMaster Campus Familiarity (%)			
Not Familiar	5%	5%	5%
Somewhat Familiar	0%	5%	15%
Neutral	41%	24%	20%
Fairly Familiar	32%	29%	25%
Very Familiar	23%	38%	35%
Orienteering Engagement (%)			

Effects of orienteering on lactate, BDNF and cognition

None	77%	90%	80%
1-2 times	18%	5%	15%
3-4 times	5%	5%	5%
Video Games (hours/week)			
None	55%	62%	55%
<1 to <3	27%	24%	30%
3 to <7	14%	10%	15%
7 to <9	5%	5%	0%
NSQ			
Egocentric	3.34 ± 0.72	3.09 ± 0.61	3.02 ± 0.84
Allocentric	3.12 ± 0.65	2.97 ± 0.61	2.85 ± 0.79
Procedural	3.65 ± 0.47	3.64 ± 0.55	3.58 ± 0.68
SAM			
Episodic	100.68 ± 15.51	100.54 ± 12.32	102.53 ± 14.07
Spatial	98.21 ± 12.91	96.10 ± 14.01	97.10 ± 12.72

351 NSQ, Navigational Strategy Questionnaire; SAM, Survey of Autobiographical memory; WC,

352 waist circumference. Values reflect M ± SD.

353

354 **Table 2. Mean Pre- and Post-Intervention Values for Primary Variables**

	Moderate Orienteering	Vigorous Orienteering	Vigorous Exercise
<i>n</i>	19	19	20
Recognition Memory			
Pre (%)	87.89 ± 10.46	87.47 ± 8.02	85.20 ± 12.67
Post (%)	84.84 ± 9.95	85.21 ± 11.57	84.15 ± 11.91
High-Interference Memory			
Pre (%)	49.89 ± 26.22	47.68 ± 17.08	45.10 ± 22.27
Post (%)	42.37 ± 30.01	49.95 ± 18.55	52.20 ± 19.82
Groton Maze Learning			

Efficiency	26.12 ± 10.32	23.28 ± 6.85	24.60 ± 5.89
Pre	29.45 ± 7.88	29.42 ± 6.45	29.92 ± 4.52
Post			
Groton Maze Test Efficiency			
Pre	35.69 ± 14.24	30.62 ± 9.69	34.19 ± 10.68
Post	38.32 ± 11.07	36.24 ± 9.61	38.42 ± 9.23
<i>n</i>	21	17	18
BDNF			
Pre BDNF (ng/mL)	40.96 ± 10.15	38.85 ± 9.30	29.00 ± 6.19
Post BDNF (ng/mL)	42.33 ± 9.75	39.78 ± 8.71	31.33 ± 5.39

355 BDNF, Brain-derived neurotrophic factor. Values reflect M ± SD.

356

357 **Intensity manipulation checks**

358 Our intervention successfully induced the appropriate level of exercise intensity for each
359 group, as confirmed by a significant main effect of group for peak lactate, $F(2, 60) = 17.49, p <$
360 $.001, \eta^2 = .37$, peak RPE $F(2, 60) = 21.56, p < .001, \eta^2 = .42$, and peak HR $F(2, 60) = 57.26, p <$
361 $.001, \eta^2 = .66$. Post hoc comparisons indicate that the moderate orienteering group had lower
362 peak HR, RPE and lactate than the vigorous orienteering and vigorous exercise groups which did
363 not differ from each other (Figure 1). Peak HR was within the instructed range of 40-50% of
364 HR_{Max} for the moderate-intensity group and 80-85% of HR_{Max} for the vigorous-intensity groups.
365 The proportion of participants above the estimated LT of 4mmol/L differed significantly between
366 groups, $H(2) = 21.70, p < .001$, with more participants above the LT in the vigorous orienteering
367 and vigorous exercise groups than the moderate orienteering group (Figure 2).

368

369 **Fig 1. Group Differences in Exercise Intensity Metrics**

370 (A) Peak HR, (B) peak RPE and (C) peak lactate achieved during the intervention between
371 groups. Bars reflect mean score, and error bars represent \pm SEM. *** = $p < .001$.

372

373 **Fig 2. Proportion of Intervention Group Above or Below Lactate Threshold**

374 Values reflect the number of participants per group with peak lactate above the lactate threshold
375 of 4mmol/L.

376

377 **Primary outcomes**

378 **BDNF**

379 Fifty-six complete cases were analyzed (moderate orienteering: $n = 21$, vigorous
380 orienteering: $n = 17$, vigorous exercise: $n = 18$). The mixed model ANOVA revealed a
381 significant main effect of time, $F(1, 53) = 10.51$, $p = .002$, $\eta_p^2 = .17$, and group, $F(2, 53) = 10.15$,
382 $p < .001$, $\eta_p^2 = .28$, but no interaction. Figure 3a shows an increase in BDNF for all groups over
383 time, but the change was only significant for the vigorous orienteering, $t(16) = 1.83$, $p = .043$, g
384 $= .42$, and the vigorous exercise groups, $t(17) = 3.09$, $p = .003$, $g = .70$, but not the moderate
385 orienteering group, $t(20) = 1.43$, $p = .084$, $g = .30$. The vigorous exercise group's BDNF levels
386 were significantly lower at baseline and post-intervention (Table 2) than the other two groups
387 (moderate orienteering, $p < .001$; vigorous orienteering, $p = .020$) (Figure 3b). This suggests that
388 the group differences seen here reflect baseline differences that are unrelated to the intervention,
389 and because of this, the relative change score (i.e., percent change) for BDNF was used in the
390 correlation and mediation analyses below.

391 Figure 4 depicts the results from the Spearman's correlation, whereby a higher peak
392 lactate achieved during exercise significantly correlated with a greater percentage increase in
393 BDNF, $r_s(54) = .28, p = .037$.

394

395 **Fig 3. Change in BDNF Concentration and Group Differences Following Intervention**

396 (A) Bars reflect mean change in BDNF concentration between groups, error bars represent \pm
397 SEM. (B) A boxplot showing the interquartile range, median, minimum, and maximum
398 concentration of BDNF between groups from pre- to post-intervention * = $p < .05$, ** = $p < .01$,
399 *** = $p < .001$.

400

401 **Fig 4. Correlation Between Peak Lactate and Percent Change in BDNF**

402 A scatterplot of ranked cases showing the correlation between the percent change in BDNF, and
403 the peak lactate achieved during exercise. $Y = 0.242x + 20.89, R = 0.28, p = .037$.

404

405 **High-interference and recognition memory**

406 Fifty-eight cases were included in the analysis (moderate orienteering: $n = 19$, vigorous
407 orienteering: $n = 19$, vigorous exercise: $n = 20$). For high-interference memory, there was a
408 significant group by time interaction, $F(1, 55) = 3.23, p = .047, \eta_p^2 = .11$. As shown in figure 5,
409 high-interference memory performance improved for the vigorous orienteering and vigorous
410 exercise groups but declined for the moderate orienteering group. The difference between the
411 moderate orienteering and vigorous exercise groups was significant, $t(37) = -2.45, p = .019, g = -$
412 $.77$. There were no other effects for high-interference memory and no effects or interaction for
413 recognition memory (Table 2).

414

415 **Fig 5. Change in High-Interference Memory Following Intervention**

416 Bars reflect mean change in performance on the lure discrimination index measure of the
417 Mnemonic Similarity Task between groups, and error bars represent \pm SEM. * = $p < .05$.

418

419 **Spatial learning and memory**

420 Sixty-one complete cases were analyzed (moderate orienteering: $n = 21$, vigorous
421 orienteering: $n = 20$, vigorous exercise: $n = 20$). Both the learning and delayed test trials of the
422 Groton Maze Learning Test revealed a significant main effect of time for both the learning
423 phase, $F(1, 58) = 30.39, p < .001, \eta_p^2 = .35$, and test phase, $F(1, 58) = 8.09, p = .006, \eta_p^2 = .12$,
424 suggesting that all groups improved in spatial processing efficiency post-intervention (Table 2).
425 For learning trials, Figure 6a depicts a significant improvement in performance for all groups
426 following the intervention, though the largest effect size was for the vigorous orienteering group,
427 $t(19) = 4.11, p < .001, g = .88$, followed by the vigorous exercise group, $t(19) = 3.43, p = .001, g$
428 $= .74$, and the moderate orienteering group, $t(20) = 2.14, p = .022, g = .45$. For the delayed test
429 performance (Figure 6b), only the vigorous orienteering group improved significantly, $t(19) =$
430 $2.70, p = .007, g = .58$. There was no effect of group or interaction for either the learning or
431 delayed test trials.

432

433 **Fig 6. Change in Spatial Learning and Memory Following Intervention**

434 (A) Bars reflect mean change in Groton Maze learning efficiency by group. (B) Bars reflect
435 mean change in Groton Maze test efficiency by group. Error bars represent \pm SEM. * = $p < .05$,

436 ** = $p < .01$, *** = $p < .001$.

437

438 **Peak lactate, BDNF and cognitive function**

439 In this exploratory analysis, the Spearman's correlation revealed significant correlations
 440 such that greater improvements in composite cognition scores were associated with higher peak
 441 lactate, $r_s(54) = .26, p = .049$, and greater increases in BDNF, $r_s(47) = .29, p = .041$. However,
 442 after controlling for the percent change in BDNF, the association between cognition and peak
 443 levels of lactate obtained during exercise was no longer significant, $r_s(44) = .21, p = .143$.

444

445 **Secondary outcomes**

446 Sixty-one complete GPS cases were analyzed (moderate orienteering, $n = 22$, vigorous
 447 orienteering, $n = 19$, vigorous exercise, $n = 20$). The mixed model ANOVA for distance travelled
 448 revealed a significant main effect of group ($F(2, 58) = 8.81, p < .001, \eta_p^2 = .23$) such that the
 449 orienteering groups travelled longer (moderate orienteering: $p = .018$; vigorous orienteering: $p <$
 450 $.001$) than those in the non-orienteering group, but the orienteering groups did not differ ($p =$
 451 $.509$). Distances travelled can be found in Table 3. Figure 7 depicts the extra distance travelled
 452 by the two orienteering groups compared to the most efficient route.

453

454 **Table 3. Distances Travelled on the Intervention Course Between Groups**

	Moderate Orienteering	Vigorous Orienteering	Vigorous Exercise
<i>n</i>	22	19	20
Average Distance Start to Midpoint (m)	678.18 ± 136.40	718.42 ± 130.74	606.5 ± 13.09

Average Distance Midpoint to Finish(m)	681.82 ± 38.62	697.37 ± 57.24	641.00 ± 13.34
Average Total Distance (m)	1360 ± 149.83*	1415.79 ± 162.25***	1247.50 ± 17.73

455 Values reflect M ± SD. * = $p < .05$ compared to the vigorous exercise group, ** = $p < .01$
 456 compared to the vigorous exercise group, *** = $p < .001$ compared to the vigorous exercise
 457 group.

458

459 **Figure 7. Routes Traversed on the Intervention Course Between Groups**

460 Figures show the routes traversed by each study group along the (approximately) 1.3 km
 461 intervention course around the McMaster University campus. Routes in pink show the paths of
 462 those in the vigorous exercise group (n = 20) who followed a researcher throughout the course at
 463 a running speed. These pink routes indicate the most efficient route. Routes in yellow show the
 464 paths of the vigorous orienteering group (n = 19) who actively navigated the intervention course
 465 at a running speed. The blue routes are those in the moderate orienteering group (n = 22) who
 466 navigated the intervention course at a walking speed. All routes were tracked using a Polar Pacer
 467 Pro GPS watch. Note that all participants started and finished in the same location, and
 468 checkpoints remained in the same location for all trials; any major differences in routes, such as
 469 a different starting location (seen in vigorous orienteering group map) can be attributed to GPS
 470 accuracy.

471

472 Across participants in the orienteering groups, those who travelled farther and, by
 473 extension, made more errors reported significantly worse egocentric spatial processing
 474 tendencies from the NSQ ($r_s(39) = -.44, p = .004$) and subjective spatial memory from the SAM
 475 ($r_s(39) = -.44, p = .004$). They also performed significantly worse on the delayed test phase of

476 the Groton Maze Test at baseline ($r_s(37) = -.35, p = .030$). Though not significant, allocentric
 477 spatial processing trended in the same direction ($r_s(39) = -.27, p = .086$). In contrast, procedural
 478 spatial processing ($r_s(39) = -.13, p = .438$) and subjective episodic memory ($r_s(39) = .05, p =$
 479 $.744$) were not related to the total distance travelled (Table 4), nor were any other aspects of
 480 cognition measured at baseline (Table 5).

481

482 **Table 4. Correlation Matrix Between Distance Travelled in the Intervention Course by the**
 483 **Orienteering Groups and Subjective Measures**

	1	2	3	4	5	6
484 1. Total Distance (m)	-					
485 2. NSQ Egocentric	-.44 **	-				
486 3. NSQ Allocentric	-.27	.55 ***	-			
487 4. NSQ Procedural	-.13	-.32 *	-.02	-		
488 5. SAM Episodic	.05	.06	-.11	-.14	-	
489 6. SAM Spatial	-.44 **	.59 ***	.49 ***	-.26	-.11	-

491 NSQ, Navigational Strategy Questionnaire; SAM, Survey of Autobiographical Memory. * =
 492 $p < .05$, ** = $p < .01$, *** = $p < .001$.

493

494 **Table 5. Correlation Matrix Between Distance Travelled in the Intervention Course by the**
 495 **Orienteering Groups and Baseline Measures of Cognitive Function**

	1	2	3	4	5
1. Total Distance (m)	-				
2. High-Interference Memory Pre	-.20				
3. Recognition Memory	-.12	.14			

496	Pre				
497	4. Groton Maze Learning	-.29	.20	.08	
498	Efficiency Pre				
499	5. Groton Maze Test	-.35*	.20	.04	.88***
500	Efficiency Pre				
501					

* = $p < .05$, *** = $p < .001$.

503

504 Discussion

505 The present study was the first to examine the effects of an acute bout of orienteering
 506 versus exercise on cognition in a sample of healthy young adults who were recreationally active
 507 but unfamiliar with orienteering. The results revealed a strong effect of exercise intensity such
 508 that the vigorous-intensity interventions in the form of either running or orienteering elicited
 509 greater increases in lactate, BDNF and memory than the moderate-intensity intervention.
 510 Additionally, vigorous orienteering improved spatial learning and memory more than vigorous
 511 running, suggesting an additional benefit of simultaneous training.

512 This study demonstrates a link between lactate, BDNF and cognition in humans. A novel
 513 and important finding is that the higher peak lactate induced by our vigorous exercise
 514 interventions was associated with greater percent increases in BDNF and better memory than our
 515 moderate-intensity intervention, lending support for the hypothesis that lactate mediates muscle-
 516 to-brain signalling [10,15,16,19]. Cognition was also significantly related to peak levels of
 517 lactate obtained during exercise. Interestingly, when controlling for BDNF, the relationship
 518 between cognition and lactate was no longer significant. We hypothesize that BDNF may partly
 519 underlie the effects of lactate on cognition, however, further work is needed to understand how
 520 exercise-induced lactate impacts cognition through and beyond its effects on BDNF
 521 [10,15,16,19].

522 On top of vigorous-intensity effects, running while navigating conferred additional
523 benefits on our measure of spatial cognition. Spatial learning and memory were tested using the
524 Groton Maze Learning Test, which is a close 2D analog to the 3D wayfinding of orienteering.
525 Although all groups increased in spatial learning efficiency, the vigorous orienteering group
526 improved the most and was the only group to improve in spatial memory after a delay. It is
527 important to consider why. One reason relates to the specific cognitive processes tested. During
528 the Groton Maze Learning Test, participants had to recall the maze route immediately and after a
529 10-minute delay, requiring skills that are highly dependent on the hippocampus, a brain region
530 that is responsive to intervention-induced plasticity [60]. A second reason why orienteering may
531 preferentially benefit spatial cognition relates to its overlap in cognitive processes engaged by
532 the task. In general, cognitive training effects tend to transfer more readily to “near-transfer”
533 tasks, i.e., tasks that closely resemble the cognitive demands of the training protocol, than “far-
534 transfer” tasks, i.e., tasks that depend on more disparate cognitive processes [61,62]. In the case
535 of orienteering, spatial cognition would classify as a near-transfer task and based on this
536 framework, would be expected to benefit the most.

537 In contrast, the high-interference memory task would be considered a far-transfer task
538 and, by the same logic, would be less likely to show additive effects, as was observed. Instead,
539 high-interference memory (lure discrimination index) improved to a similar extent for both
540 vigorous exercise and orienteering groups, suggesting that this aspect of cognition is more
541 sensitive to the acute effects of exercise intensity than the combined effects of the exercise-
542 cognitive training that is experienced during an acute bout of orienteering. Although the effects
543 of vigorous exercise on high-interference memory were expected and consistent with prior work
544 [9,56,57,63], we were surprised to observe a decrement in high-interference memory

545 performance following moderate-intensity orienteering. This may be related to the amount of
546 exercise-induced BDNF, which is less after moderate intensities compared to vigorous [6–8].
547 Indeed, those who orienteered at moderate intensity produced less BDNF than those who
548 orienteered at a vigorous intensity, and this may have reduced their neurogenic support,
549 rendering substrate-dependent memory benefits unobtainable.

550 The difference in BDNF levels between moderate and vigorous intensity orienteering
551 may also help to explain why expert taxi drivers experience a trade-off that augments their
552 posterior (primarily relating to spatial processing) hippocampus at the cost of their anterior
553 (mainly involved in episodic memory) hippocampus [39]. Taxi drivers are sedentary while
554 driving, which is in stark contrast to expert orienteers who perform their sport at a rapid running
555 speed [29]. The lack of vigorous movement during navigation may be why we see evidence for a
556 trade-off in expert taxi drivers but not in expert orienteers. Regardless of the mechanism, we
557 found that engaging in vigorous-intensity exercise while orienteering benefited spatial memory
558 but not high-interference memory. Although the reasons for this are unclear, a single acute
559 orienteering session may not be a strong enough stimulus to evoke adaptative changes in all
560 hippocampal functions. Future research is needed to investigate whether chronic orienteering
561 interventions can produce “far-transfer” effects beyond the effect of spatial cognition observed
562 here.

563 Although BDNF increased more for the vigorous interventions than the moderate
564 intervention, it was expected to increase even more following vigorous orienteering, but that was
565 not observed. Our sample of healthy, recreationally active younger adults may have been the
566 reason why. Unlike older adults, prior research with younger adults reveals no additional boost in
567 BDNF from exercise-cognitive training, as was observed here [57,64]. This makes sense given

568 that BDNF is thought to respond to energetic challenges [65], and in our sample of recreationally
569 active younger adults, the additional challenge of running while navigating may not have been
570 enough of an acute energetic demand. This may be especially true given the wayfinding task was
571 short (only ~12 minutes) and across a familiar terrain. Future work should examine the
572 potentially additive effects of orienteering versus running on BDNF using longer and less
573 familiar routes. Additionally, there is evidence that females have lower BDNF responsivity to
574 acute exercise [66] and lower lactate responses at the same relative exercise intensity compared
575 to males [67]. These potential sex-based differences in lactate-induced BDNF activation may be
576 at play with our predominately female sample (65%) and should be followed up in future work.

577 Despite our participants' familiarity with the campus on which the orienteering course
578 was set, both the moderate and vigorous intensity orienteering groups travelled significantly
579 farther and, by extension, made more errors than the most efficient route. Interestingly, the
580 distance travelled while orienteering was associated with several of our baseline measures.
581 Notably, those who travelled shorter distances (i.e., made fewer errors) had better spatial
582 memory at baseline, as revealed by both self-report and task performance, which reaffirms the
583 existence of overlapping cognitive processes engaged between navigation and spatial memory
584 [30]. Also, those who travelled shorter distances reported greater reliance on egocentric spatial
585 navigation. Allocentric spatial navigation was not as strongly related to course distance travelled,
586 which was surprising given that allocentric spatial processing, like egocentric spatial processing
587 and spatial memory, have been previously associated with expertise in the sport of orienteering
588 [27]. The weak association between allocentric spatial processing and navigational efficiency
589 observed here may be related to participants' familiarity with the course terrain. We set the
590 course on campus because it provided a safe environment for orienteering, but it is important to

591 acknowledge that navigational tendencies may differ between familiar and unfamiliar terrains
592 [30]. For example, participants could identify campus buildings by their names and then navigate
593 based on previously learned routes rather than utilize allocentric spatial navigation. It will be
594 important for future work to examine the orienteering interventions across novel terrains over a
595 variety of course difficulties.

596 Moreover, overreliance on GPS devices may be a factor because it minimizes active
597 navigation and the practice of allocentric navigation in the case of “use it or lose it” [36]. GPS
598 may be used more commonly by those with little experience in orienteering, as allocentric
599 navigation may require more practice to be developed [30]. Unfortunately, we did not capture
600 GPS use, but we would recommend this be done in future studies. Furthermore, prior research
601 suggests that females may rely less on allocentric navigation than males [68], and our sample
602 was predominantly female. Although we did not power our sample size to examine sex
603 differences, it is recommended that future research do so. Another reason why we failed to
604 observe a strong association between allocentric spatial processing and navigational efficiency is
605 speculative but worth noting; this study was conducted in North America where orienteering
606 awareness and practice is relatively limited compared to Nordic countries where orienteering is
607 embedded into the school curricula and local cultural activities [69]. This fact should be
608 considered when comparing studies from different countries.

609

610 **Conclusion**

611 This study demonstrates the effect of vigorous exercise on lactate, BDNF and
612 hippocampal-dependent memory. It also reveals that orienteering may outperform exercise in
613 improving spatial memory when done at a vigorous intensity. Together, this study establishes the

614 efficacy of using orienteering to improve cognition in younger adults and provides essential
615 groundwork for future research in older adult or AD populations to help preserve cognitive
616 function across the lifespan.

617

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622

623 **References**

- 624 1. Burke SN, Barnes CA. Neural plasticity in the ageing brain. *Nat Rev Neurosci*. 2006 Jan
625 1;7(1):30–40.
- 626 2. Fraser MA, Shaw ME, Cherbuin N. A systematic review and meta-analysis of longitudinal
627 hippocampal atrophy in healthy human ageing. *NeuroImage*. 2015 May;112:364–74.
- 628 3. Jack CR, Petersen RC, Xu Y, O’Brien PC, Smith GE, Ivnik RJ, et al. Rate of medial temporal
629 lobe atrophy in typical aging and Alzheimer’s disease. *Neurology*. 1998 Oct;51(4):993–9.
- 630 4. Deweer B, Lehericy S, Pillon B, Baulac M, Chiras J, Marsault C, et al. Memory disorders in
631 probable Alzheimer’s disease: the role of hippocampal atrophy as shown with MRI. *J*
632 *Neurol Neurosurg Psychiatry*. 1995 May 1;58(5):590–7.
- 633 5. Nichols E, Steinmetz JD, Vollset SE, Fukutaki K, Chalek J, Abd-Allah F, et al. Estimation of
634 the global prevalence of dementia in 2019 and forecasted prevalence in 2050: an analysis
635 for the Global Burden of Disease Study 2019. *Lancet Public Health*. 2022 Feb;7(2):e105–
636 25.
- 637 6. Griffin ÉW, Mullally S, Foley C, Warmington SA, O’Mara SM, Kelly ÁM. Aerobic exercise
638 improves hippocampal function and increases BDNF in the serum of young adult males.
639 *Physiol Behav*. 2011 Oct;104(5):934–41.
- 640 7. Shah Z, Ahmad F, Zahra M, Zulfiqar F, Aziz S, Mahmood A. Effect of Single Bout of
641 Moderate and High Intensity Interval Exercise on Brain Derived Neurotrophic Factor and
642 Working Memory in Young Adult Females. van Praag H, Wrann C, editors. *Brain Plast*.
643 2022 Oct 21;8(1):35–42.

- 644 8. Jeon YK, Ha CH. The effect of exercise intensity on brain derived neurotrophic factor and
645 memory in adolescents. *Environ Health Prev Med.* 2017 Dec;22(1):27.
- 646 9. Kovacevic A, Fenesi B, Paolucci E, Heisz JJ. The effects of aerobic exercise intensity on
647 memory in older adults. *Appl Physiol Nutr Metab.* 2020 Jun;45(6):591–600.
- 648 10. Hashimoto T, Tsukamoto H, Takenaka S, Olesen ND, Petersen LG, Sørensen H, et al.
649 Maintained exercise-enhanced brain executive function related to cerebral lactate
650 metabolism in men. *FASEB J.* 2018;32(3):1417–27.
- 651 11. Faude O, Kindermann W, Meyer T. Lactate Threshold Concepts: How Valid are They?
652 *Sports Med.* 2009 May;39(6):469–90.
- 653 12. Heck H, Mader A, Hess G, Mücke S, Müller R, Hollmann W. Justification of the 4-mmol/l
654 Lactate Threshold. *Int J Sports Med.* 1985 Jun;06(03):117–30.
- 655 13. Passarella S, de Bari L, Valenti D, Pizzuto R, Paventi G, Atlante A. Mitochondria and L -
656 lactate metabolism. *FEBS Lett.* 2008 Oct 29;582(25–26):3569–76.
- 657 14. Ide K, Schmalbruch IK, Quistorff B, Horn A, Secher NH. Lactate, glucose and O₂ uptake
658 in human brain during recovery from maximal exercise. *J Physiol.* 2000;522(1):159–64.
- 659 15. El Hayek L, Khalifeh M, Zibara V, Abi Assaad R, Emmanuel N, Karnib N, et al. Lactate
660 mediates the effects of exercise on learning and memory through SIRT1-dependent
661 activation of hippocampal brain-derived neurotrophic factor (BDNF). *J Neurosci.* 2019 Jan
662 28;1661–18.
- 663 16. Xue X, Liu B, Hu J, Bian X, Lou S. The potential mechanisms of lactate in mediating
664 exercise-enhanced cognitive function: a dual role as an energy supply substrate and a
665 signaling molecule. *Nutr Metab.* 2022 Dec;19(1):52.
- 666 17. van Hall G, Stømstad M, Rasmussen P, Jans Ø, Zaar M, Gam C, et al. Blood Lactate is an
667 Important Energy Source for the Human Brain. *J Cereb Blood Flow Metab.* 2009
668 Jun;29(6):1121–9.
- 669 18. Hashimoto T, Tsukamoto H, Ando S, Ogoh S. Effect of Exercise on Brain Health: The
670 Potential Role of Lactate as a Myokine. *Metabolites.* 2021 Nov 29;11(12):813.
- 671 19. Müller P, Duderstadt Y, Lessmann V, Müller NG. Lactate and BDNF: Key Mediators of
672 Exercise Induced Neuroplasticity? *J Clin Med.* 2020 Apr;9(4):1136.
- 673 20. Chang YK, Labban JD, Gapin JI, Etnier JL. The effects of acute exercise on cognitive
674 performance: A meta-analysis. *Brain Res.* 2012 May;1453:87–101.
- 675 21. Kujach S, Olek RA, Byun K, Suwabe K, Sitek EJ, Ziemann E, et al. Acute Sprint Interval
676 Exercise Increases Both Cognitive Functions and Peripheral Neurotrophic Factors in
677 Humans: The Possible Involvement of Lactate. *Front Neurosci.* 2020 Jan 23;13:1455.

- 678 22. Tsukamoto H, Suga T, Takenaka S, Tanaka D, Takeuchi T, Hamaoka T, et al. Repeated
679 high-intensity interval exercise shortens the positive effect on executive function during
680 post-exercise recovery in healthy young males. *Physiol Behav.* 2016 Jun;160:26–34.
- 681 23. Tsukamoto H, Suga T, Takenaka S, Tanaka D, Takeuchi T, Hamaoka T, et al. Greater
682 impact of acute high-intensity interval exercise on post-exercise executive function
683 compared to moderate-intensity continuous exercise. *Physiol Behav.* 2016 Mar;155:224–
684 30.
- 685 24. Kronenberg G, Reuter K, Steiner B, Brandt MD, Jessberger S, Yamaguchi M, et al.
686 Subpopulations of proliferating cells of the adult hippocampus respond differently to
687 physiologic neurogenic stimuli. *J Comp Neurol.* 2003 Dec 22;467(4):455–63.
- 688 25. Gavelin HM, Dong C, Minkov R, Bahar-Fuchs A, Ellis KA, Lautenschlager NT, et al.
689 Combined physical and cognitive training for older adults with and without cognitive
690 impairment: A systematic review and network meta-analysis of randomized controlled
691 trials. *Ageing Res Rev.* 2021 Mar;66:101232.
- 692 26. Lövdén M, Schaefer S, Noack H, Bodammer NC, Kühn S, Heinze HJ, et al. Spatial
693 navigation training protects the hippocampus against age-related changes during early and
694 late adulthood. *Neurobiol Aging.* 2012 Mar;33(3):620.e9-620.e22.
- 695 27. Waddington EE, Heisz JJ. Orienteering experts report more proficient spatial processing
696 and memory across adulthood. Kavushansky A, editor. *PLOS ONE.* 2023 Jan
697 20;18(1):e0280435.
- 698 28. Eccles DW, Walsh SE, Ingledew DK. A grounded theory of expert cognition in
699 orienteering. *J Sport Exerc Psychology.* 2002;24:68–88.
- 700 29. Eccles DW, Walsh SE, Ingledew DK. Visual attention in orienteers at different levels of
701 experience. *J Sports Sci.* 2006 Jan;24(1):77–87.
- 702 30. Ekstrom AD, Hill PF. Spatial navigation and memory: A review of the similarities and
703 differences relevant to brain models and age. *Neuron.* 2023 Apr;111(7):1037–49.
- 704 31. Nedelska Z, Andel R, Laczó J, Vlcek K, Horinek D, Lisy J, et al. Spatial navigation
705 impairment is proportional to right hippocampal volume. *Proc Natl Acad Sci.* 2012 Feb
706 14;109(7):2590–4.
- 707 32. Serino S, Giuseppe R. What is the role of spatial processing in the decline of episodic
708 memory in Alzheimer’s disease? The “mental frame syncing” hypothesis. 2014;
- 709 33. Boccia M, Di Vita A, Diana S, Margiotta R, Imbriano L, Rendace L, et al. Is Losing One’s
710 Way a Sign of Cognitive Decay? Topographical Memory Deficit as an Early Marker of
711 Pathological Aging. *J Alzheimers Dis.* 2019 Mar 29;68(2):679–93.
- 712 34. Moffat SD, Resnick SM. Effects of age on virtual environment place navigation and
713 allocentric cognitive mapping. *Behav Neurosci.* 2002 Oct;116(5):851–9.

- 714 35. McKinlay R. Technology: Use or lose our navigation skills. *Nature*. 2016 Mar
715 31;531(7596):573–5.
- 716 36. Dahmani L, Bohbot VD. Habitual use of GPS negatively impacts spatial memory during
717 self-guided navigation. *Sci Rep*. 2020 Dec;10(1):6310.
- 718 37. Ishikawa T. Satellite Navigation and Geospatial Awareness: Long-Term Effects of Using
719 Navigation Tools on Wayfinding and Spatial Orientation. *Prof Geogr*. 2019 Apr
720 3;71(2):197–209.
- 721 38. Wilson M. Six views of embodied cognition. *Psychon Bull Rev*. 2002 Dec;9(4):625–36.
- 722 39. Maguire EA, Gadian DG, Johnsrude IS, Good CD, Ashburner J, Frackowiak RSJ, et al.
723 Navigation-related structural change in the hippocampi of taxi drivers. *Proc Natl Acad Sci*.
724 2000 Apr 11;97(8):4398–403.
- 725 40. Sheldon S, Levine B. The role of the hippocampus in memory and mental construction:
726 Memory and mental construction. *Ann N Y Acad Sci*. 2016 Apr;1369(1):76–92.
- 727 41. Vogel JW, La Joie R, Grothe MJ, Diaz-Papkovich A, Doyle A, Vachon-Preseu E, et al. A
728 molecular gradient along the longitudinal axis of the human hippocampus informs large-
729 scale behavioral systems. *Nat Commun*. 2020 Feb 19;11(1):960.
- 730 42. Bao S, Liu J, Liu Y. Shedding Light on the Effects of Orienteering Exercise on Spatial
731 Memory Performance in College Students of Different Genders: An fNIRS Study. *Brain*
732 *Sci*. 2022 Jun 29;12(7):852.
- 733 43. Feraco T, Bonvento M, Meneghetti C. Orienteering: What relation with visuospatial
734 abilities, wayfinding attitudes, and environment learning? *Appl Cogn Psychol*. 2021
735 Nov;35(6):1592–9.
- 736 44. Roca-González C, Martín Gutierrez J, García-Dominguez M, Mato Carrodegua M del C.
737 Virtual Technologies to Develop Visual-Spatial Ability in Engineering Students. *EURASIA*
738 *J Math Sci Technol Educ [Internet]*. 2016 Dec 2 [cited 2022 Nov 8];13(2). Available from:
739 [https://www.ejmste.com/article/virtual-technologies-to-develop-visual-spatial-ability-in-](https://www.ejmste.com/article/virtual-technologies-to-develop-visual-spatial-ability-in-engineering-students-4673)
740 [engineering-students-4673](https://www.ejmste.com/article/virtual-technologies-to-develop-visual-spatial-ability-in-engineering-students-4673)
- 741 45. Canadian Society for Exercise Physiology. Canadian Society for Exercise Physiology-
742 Physical Activity Training for Health (CSEP-PATH). 2013;
- 743 46. Mottet M, Saury J. Accurately locating one’s spatial position in one’s environment during a
744 navigation task: Adaptive activity for finding or setting control flags in orienteering.
745 *Psychol Sport Exerc*. 2013 Mar;14(2):189–99.
- 746 47. Zhong JY, Kozhevnikov M. Relating allocentric and egocentric survey-based
747 representations to the self-reported use of a navigation strategy of egocentric spatial
748 updating. *J Environ Psychol*. 2016;46:154–75.

- 749 48. Palombo DJ, Williams LJ, Abdi H, Levine B. The survey of autobiographical memory
750 (SAM): A novel measure of trait mnemonics in everyday life. *Cortex*. 2013
751 Jun;49(6):1526–40.
- 752 49. Tanaka H, Monahan KD, Seals DR. Age-predicted maximal heart rate revisited. *J Am Coll*
753 *Cardiol*. 2001 Jan;37(1):153–6.
- 754 50. Nes BM, Janszky I, Vatten LJ, Nilsen TIL, Aspenes ST, Wisløff U. Estimating V̇O₂peak
755 from a Nonexercise Prediction Model: The HUNT Study, Norway. *Med Sci Sports Exerc*.
756 2011 Nov;43(11):2024–30.
- 757 51. Borg G AV. Perceived Exertion and Pain Scales. *Medicine & science in sports & exercise*.
758 1982;
- 759 52. Bakker A, Kirwan CB, Miller M, Stark CEL. Pattern Separation in the Human
760 Hippocampal CA3 and Dentate Gyrus. *Science*. 2008 Mar 21;319(5870):1640–2.
- 761 53. Kirwan CB, Stark CEL. Overcoming interference: An fMRI investigation of pattern
762 separation in the medial temporal lobe. *Learn Mem*. 2007 Sep;14(9):625–33.
- 763 54. Yassa MA, Stark CEL. Pattern separation in the hippocampus. *Trends Neurosci*. 2011
764 Oct;34(10):515–25.
- 765 55. Stark SM, Kirwan CB, Stark CEL. Mnemonic Similarity Task: A Tool for Assessing
766 Hippocampal Integrity. *Trends Cogn Sci*. 2019 Nov;23(11):938–51.
- 767 56. Déry N, Pilgrim M, Gibala M, Gillen J, Wojtowicz JM, MacQueen G, et al. Adult
768 hippocampal neurogenesis reduces memory interference in humans: opposing effects of
769 aerobic exercise and depression. *Front Neurosci* [Internet]. 2013 [cited 2023 Aug 2];7.
770 Available from: <http://journal.frontiersin.org/article/10.3389/fnins.2013.00066/abstract>
- 771 57. Heisz JJ, Clark IB, Bonin K, Paolucci EM, Michalski B, Becker S, et al. The Effects of
772 Physical Exercise and Cognitive Training on Memory and Neurotrophic Factors. *J Cogn*
773 *Neurosci*. 2017 Nov;29(11):1895–907.
- 774 58. Milner B. Some effects of frontal lobectomy in man. The frontal granular cortex and
775 behavior. 1964;313–34.
- 776 59. Pietrzak R, Cohen H, Snyder P. Spatial learning efficiency and error monitoring in normal
777 aging: An investigation using a novel hidden maze learning test. *Arch Clin Neuropsychol*.
778 2007 Feb;22(2):235–45.
- 779 60. Byrne P, Becker S, Burgess N. Remembering the past and imagining the future: A neural
780 model of spatial memory and imagery. *Psychol Rev*. 2007 Apr;114(2):340–75.
- 781 61. Sala G, Aksayli ND, Tatlidil KS, Tatsumi T, Gondo Y, Gobet F. Near and Far Transfer in
782 Cognitive Training: A Second-Order Meta-Analysis. Zwaan R, Verkoeijen P, editors.
783 *Collabra Psychol*. 2019;5(1):18.

- 784 62. Sala G, Aksayli ND, Tatlidil KS, Gondo Y, Gobet F. Working memory training does not
785 enhance older adults' cognitive skills: A comprehensive meta-analysis. *Intelligence*.
786 2019;77:101386.
- 787 63. Crawford L, Loprinzi P. Effects of Intensity-Specific Acute Exercise on Paired-Associative
788 Memory and Memory Interference. *Psych*. 2019 May 25;1(1):290–305.
- 789 64. Miyamoto T, Hashimoto S, Yanamoto H, Ikawa M, Nakano Y, Sekiyama T, et al. Response
790 of brain-derived neurotrophic factor to combining cognitive and physical exercise. *Eur J*
791 *Sport Sci*. 2018 Sep 14;18(8):1119–27.
- 792 65. Marosi K, Mattson MP. BDNF mediates adaptive brain and body responses to energetic
793 challenges. *Trends Endocrinol Metab*. 2014 Feb;25(2):89–98.
- 794 66. Dinoff A, Herrmann N, Swardfager W, Lanctôt KL. The effect of acute exercise on blood
795 concentrations of brain-derived neurotrophic factor in healthy adults: a meta-analysis. Foxe
796 J, editor. *Eur J Neurosci*. 2017 Jul;46(1):1635–46.
- 797 67. Wheatley CM, Snyder EM, Johnson BD, Olson TP. Sex differences in cardiovascular
798 function during submaximal exercise in humans. *SpringerPlus*. 2014 Dec;3(1):445.
- 799 68. Grön G, Wunderlich AP, Spitzer M, Tomczak R, Riepe MW. Brain activation during
800 human navigation: gender-different neural networks as substrate of performance. *Nat*
801 *Neurosci*. 2000 Apr;3(4):404–8.
- 802 69. Coutrot A, Silva R, Manley E, de Cothi W, Sami S, Bohbot VD, et al. Global Determinants
803 of Navigation Ability. *Curr Biol*. 2018 Sep;28(17):2861-2866.e4.

804

805 **Supporting Information**

806

807 **S1 Appendix. Demographic Questionnaire.** The included questions comprised the
808 demographics questionnaire administered in the online baseline questionnaire.