



CREaTE

Canterbury Research and Theses Environment

Canterbury Christ Church University's repository of research outputs

<http://create.canterbury.ac.uk>

Please cite this publication as follows:

Hurst, P., Foad, A., Coleman, D. A. and Beedie, C. (2017) Athletes intending to use sports supplements are more likely to respond to a placebo. *Medicine & Science in Sports & Exercise* (MSSE). ISSN 0195-9131.

Link to official URL (if available):

<http://dx.doi.org/10.1249/MSS.0000000000001297>

This version is made available in accordance with publishers' policies. All material made available by CReaTE is protected by intellectual property law, including copyright law. Any use made of the contents should comply with the relevant law.

Contact: create.library@canterbury.ac.uk



1 **Athletes Intending to Use Sports Supplements Are More Likely to Respond to a Placebo**

2 Philip Hurst¹, Abby Foad¹, Damian Coleman¹ and Chris Beedie¹

3 ¹Canterbury Christ Church University

4 Address for Correspondence: Philip Hurst, Canterbury Christ Church University, North

5 Holmes Road, Canterbury, CT1 1QU

6 Telephone: 01227 923130

7 Email: philip.hurst@canterbury.ac.uk

8 **Abstract**

9 **Purpose:** We investigated associations between athletes' use of sport supplements and their
10 responsiveness to placebo and nocebo interventions. **Methods:** Participants (n=627) reported
11 their intention to use, and actual use of, sport supplements. They then completed a 5x20m
12 repeat sprint protocol in the baseline condition, prior to being randomized to one of three
13 treatments. Participants in the positive-belief treatment were administered an inert capsule
14 described as a potent supplement which would improve sprint performance. Participants in the
15 negative-belief treatment were administered an inert capsule described as a potent supplement
16 which would negatively affect sprint performance. Participants in the control treatment
17 received neither instruction nor capsule. 20 minutes following baseline trials, all participants
18 completed the same repeat sprint protocol in the experimental condition. **Results:** Compared
19 to controls, no mean differences in performance were observed between baseline and
20 experimental conditions for the positive-belief treatment ($-0.07 \pm 0.27\%$, $d=0.02$), but mean
21 differences were observed for the negative-belief treatment ($-0.92 \pm 0.31\%$, $d=0.32$),
22 suggesting a moderate nocebo effect. In the positive-belief treatment however, a relationship
23 between intention to use supplements and performance was observed. Performance worsened
24 by $-1.10\% \pm 0.30\%$ compared to baseline for participants not intending to use supplements,
25 worsened by $-0.64 \pm 0.43\%$ among those undecided about supplement use, but improved by
26 $0.19 \pm 0.24\%$ among those participants intending to use supplements. **Conclusion:** Information
27 about a harmful supplement worsened repeat sprint performance (a mean nocebo effect),
28 whereas information about a beneficial supplement did not improve performance (no mean
29 placebo effect was observed). However, participants' intention to use sport supplements
30 influenced the direction and magnitude of subsequent placebo responses, with participants
31 intending to use supplements more likely to respond to the positive intervention.

32 **Key words:** Nocebo, responders, beliefs, ergogenic aids

33 **Introduction**

34 A placebo effect is a positive psychobiological response to a purported beneficial treatment
35 (11). Placebo effects have been extensively studied in sport (3, 4, 7, 8, 12, 15, 21, 31, 32, 34,
36 35, 39), with a systematic review (6) reporting that placebo treatments can exert a significant
37 effect on sport performance. For example, Ross et al. (34) reported a 1.2% improvement in 3-
38 km running time-trial performance when participants self-administered saline injections
39 believing it to be a performance enhancing substance. Likewise, Saunders et al. (35) reported
40 that mean power output improved by 3.7% among cyclists deceptively administered a placebo
41 when they believed they had ingested caffeine.

42 While there is empirical support for the potential role of the placebo effect in sports
43 performance, there is less evidence for the nocebo effect; that is, a negative psychobiological
44 response to a purported harmful treatment. Arguably the first study of the nocebo effect in sport
45 was conducted by Beedie et al. in 2007 (5). These authors reported that n=21 participants who
46 believed they had ingested a placebo, that is a capsule described as a beneficial sport
47 supplement, ran progressively faster compared to baseline. Likewise, n=21 participants who
48 believed they had ingested a nocebo, that is a capsule described as a supplement likely to be
49 detrimental to performance, ran progressively slower compared to baseline. Findings
50 highlighted the potentially significant impact of positive and negative expectations on sports
51 performance.

52 However, the study in question (5) lacked a no-treatment control. It is therefore problematic to
53 estimate the true relative magnitude of the placebo and nocebo effects reported; changes in
54 performance could be attributed to statistical or methodological artefacts such as regression to
55 the mean or spontaneous improvements/decrements in performance (25). Further, it is
56 problematic from this uncontrolled study to discern whether actual effects were all positive, all
57 negative, or whether both placebo and nocebo effects occurred. As a result, the reported

58 magnitude of either the nocebo or placebo effect might have been overestimated. Further, while
59 the n=42 reported was relatively large for an intervention study in sport, it was however too
60 small to facilitate the reliable identification of any psychosocial variables that might have been
61 associated with the placebo and nocebo responses observed.

62 In most studies of the placebo/nocebo effect in sport, the standard deviation of the dependent
63 measure is greater in experimental conditions than at baseline (6). This suggests that, even if a
64 mean placebo effect is observed, there is considerable inter-individual variability in response
65 to the treatment. Few studies have attempted to identify the variables related to placebo
66 responses, and those that have are perhaps methodologically unsatisfactory. For example,
67 Beedie et al. (7) identified a possible link between placebo responding and personality factors,
68 but the sample size was too small for their findings to be considered reliable. In fact, the small
69 sample sizes of nearly all studies of the placebo effect in sport has precluded the reliable
70 investigation of any factor that might be associated with placebo responding. If our knowledge
71 and understanding of the placebo and nocebo effects is to progress beyond simple description,
72 we need to better understand the relevant antecedents and mechanisms.

73 We aimed to extend Beedie et al.'s study (5) via two specific criteria, each allowing us to test
74 two novel hypotheses. First, by using a no-treatment control we were able to estimate the
75 relative magnitude of placebo and nocebo effects in response to treatments. In this context we
76 hypothesised that compared to controls, positive effects on performance would be associated
77 with a positive belief (placebo) treatment, while negative effects on performance would be
78 associated with a negative belief (nocebo) treatment. Second, by using a sufficiently large
79 sample, we were able to reliably identify factors that might be associated with observed placebo
80 and/or nocebo responses. Given the range of such factors is potentially large, we were
81 presented with a number of possible hypotheses. Recent data from both medicine and
82 psychology suggest that prior use of a treatment can influence the response of a patient to a

83 subsequent placebo treatment (10). We hypothesised that athletes with prior experience of sport
84 supplements would be more likely to respond to a placebo sport supplement than those who do
85 not use sport supplements. Furthermore, prior use of a supplement is suggested to be influenced
86 via a person's intention to use that substance (33). We therefore further hypothesised that those
87 intending to use supplements would also be more likely to respond to a placebo intervention.

88 The idea that greater understanding of the placebo effect among athletes and coaches might
89 reduce doping has been proposed (6, 26, 31, 32). Given the gateway hypothesis (26), which
90 posits that supplement use can lead to doping, it is reasonable to suggest that, over and above
91 enhancing our understanding of placebo and nocebo effects in sport, this study could also
92 enhance our understanding of factors that underpin doping.

93 **Methods**

94 Design

95 The placebo and nocebo interventions used in this study required the deceptive administration
96 of an inert capsule delivered to members of teams in their usual team environment. We
97 therefore used a cluster randomized controlled trial design to minimize cross-contamination
98 between experimental and control treatments. Participants completed a pre-experimental
99 questionnaire relating to sport supplementation, before performing 5 × 20-m repeat sprint with
100 30s recovery at baseline. Following Beedie et al.'s original design (5), participants in the
101 positive-belief treatment (n = 288) were deceptively administered an inert capsule described as
102 a potent supplement which would improve sprint performance. Also following the original
103 design, participants in the negative-belief treatment (n = 232) were deceptively administered
104 an inert capsule described as a potent supplement which would negatively affect sprint
105 performance. However, extending the original study, no-treatment control participants (n =
106 192) received neither instruction nor placebo. Twenty minutes following the administration of

107 the capsules, participants completed the experimental condition, which was a repeat of the 5 ×
108 20-m sprints.

109 Participants

110 We used convenience sampling, and invited athletes from a range of sports to participate in the
111 study. Seven hundred and twelve competitive athletes from 43 different teams (number of
112 athletes in each team: median = 14; range = 8 to 40) were initially recruited to the study.
113 Participant demographics are presented in Table 1. All participants were aware that their
114 involvement in the study was voluntary and that all data collected would be treated as
115 confidential. Ethical approval was granted by the Institutional Research Ethics Committee.
116 Participants gave written informed consent once they had read the participant information
117 sheet.

118 Measures

119 Pre-experimental questionnaire

120 All participants were asked to complete a pre-experimental questionnaire detailing sex, age,
121 sport played and competitive level (club, county, regional or national). They were asked to
122 indicate whether they used sports supplements (yes or no), the total number of supplements
123 used, and the frequency of use (daily, weekly, monthly or never). They were also asked to
124 indicate their agreement with a statement of their intention to use sport supplements in the next
125 three months on a 6 point Likert-type scale anchored at strongly disagree (1), through to
126 strongly agree (6). Those scoring 1 and 2 were grouped as ‘not intending’, 3 and 4 as
127 ‘undecided’, 5 and 6 as ‘intending’.

128 Repeat sprint performance

129 Whereas Beedie et al. (5) used a 3 × 30-m repeat sprint protocol, Schimpchen, Skorski, Nopp
130 and Meyer (36) reported that four or more sprints should be used to decrease the typical error

131 and improve the precision of estimating true changes in performance. Furthermore, the
132 majority of sprinting in team sports events occurs over relatively short distances (i.e. <30-m;
133 (14)) and short durations (i.e. <4 seconds; (37)). For these reasons, participants were asked to
134 complete five 20-m maximal intensity repeat sprints with 30 seconds of recovery between each
135 sprint. Sprint time was measured using an automated, single-beam photocell, light gate system
136 (Smartspeed ProTM, Fusion Sport Inc., Australia). Single-beam light gate systems are the most
137 common method for measuring sprint performance and have been shown to have good
138 reliability (20).

139 Belief Manipulation

140 During the 20-minute recovery period between baseline and experimental conditions,
141 participants in the positive- and negative-belief treatments were given a capsule described as a
142 potent sport supplement, ‘inorganic nitrate.’ Similar to Beedie et al. (5), the positive-belief
143 treatment participants were given two red and white, size 1 (20-mm), gelatine capsules
144 containing 200-mg of cornflour (Sainsbury’s, London UK) and informed that inorganic nitrate
145 would improve both endurance and repeat sprint performance. Negative-belief treatment
146 participants were given two red and black, size 1 (20-mm), gelatine capsules containing 200-
147 mg of cornflour and informed that inorganic nitrate would improve endurance but have a
148 negative effect on sprint speed. The effectiveness of the belief manipulation was assessed
149 during a debrief immediately following the experimental trials, at which point the true nature
150 of the study was revealed. Participants were asked to respond on a 10 point Likert-type scale,
151 how much they believed the treatment influenced their performance (1 = no influence to 10 =
152 high influence).

153 Procedure

154 Testing was performed at the 43 different training facilities habitually used by the teams
155 recruited to the study. All data per each participant were collected on one day to minimize
156 meteorological and biological variation. Teams were randomised to the three treatments (i.e.
157 positive, negative and control) using a computer generated cluster programme (allocation ratio
158 1:1:1), which was performed by the lead author who was also involved in delivering the
159 intervention. To reduce potential confounding, only one team per club were permitted to take
160 part in the study. All treatments were conducted on separate days and at separate sites to
161 maintain the experimental blind.

162 Participants completed the sprints in footwear and clothing suitable for high intensity exercise,
163 and were encouraged to perform their standard warm-up. They began each sprint in a stationary
164 position, ~50-cm behind the first light gate. They were instructed not to rock back and forth
165 prior to the sprint, but were permitted to start the sprint in any position (e.g. split-stance or
166 crouch start), which was replicated for each sprint. Each sprint was started by a green LED,
167 which would flash up on the photocell. Participants were encouraged to sprint as fast as
168 possible for the full 20-m, with times recorded to the nearest 1/100th of a second. Participants
169 were given thirty seconds to jog back to the start position and begin the next sprint. This process
170 was continued until each participant had completed five sprints.

171 After the baseline condition, participants in the positive- and negative-belief treatments
172 received the capsules and the belief manipulation. All participants then completed a 20-minute
173 recovery consisting of light exercise to minimize the search for physiological symptoms
174 associated with the intervention (16), before commencing the experimental condition in the
175 same manner as the first. The total duration of the repeat sprint protocol, including recovery,
176 was less than 30-minutes per participant. On completion, participants were debriefed about the
177 true nature of the study in line with American Psychological Association guidelines for
178 deceptive research (1).

179 Statistical analysis

180 Data were inputted into SPSS version 23.0 (IBM, Armonk, NY, USA) and tested for
181 homogeneity of variance, normal distribution and anomalies. Inspection of the data indicated
182 that 55 participants (8%) did not complete the experimental condition (positive-belief treatment
183 n = 20; negative-belief treatment n = 16; control n = 19). In addition, data values that exceeded
184 2.5 times the standard deviation were identified as extreme outliers (30). Thirty participants
185 (4%) were identified as extreme outliers (positive-belief treatment n = 7; negative-belief
186 treatment n = 7; control n = 16) and were subsequently removed from further analysis (27).
187 Data for the remaining sample of 627 participants (positive-belief treatment n = 261; negative-
188 belief treatment n = 209; control n = 157) were entered into subsequent statistical analyses.

189 One-way Analysis of Variance (ANOVA) and chi-square (χ^2) tests were used to compare
190 continuous (years training, hours per week training and number of supplement used) and
191 categorical (sex, age, sport, ability, supplement use, frequency of supplement use and intention
192 to use supplements) variables between treatments, respectively.

193 Sprint times for each condition (i.e. baseline and experimental) and treatment (i.e. positive,
194 negative and control) were inputted into Hopkins' (22) reliability spreadsheet. Data were log
195 transformed to reduce non-uniform errors and the intra-class correlation (ICC) provided
196 estimates of reliability. The precision of ICC was interpreted as extremely high = 0.99; very
197 high = 0.90; high = 0.75; moderate = 0.50; low = 0.20 (22).

198 Hopkins, Hawley and Burke (24) suggest that research investigating athletic performance
199 should report outcome as a percentage change from baseline. Sprint times were therefore
200 converted to the proportion of the first sprint speed, expressed as a percentage. Differences
201 between participant's average performance for each condition (i.e. performance average for
202 baseline [sprints 1 to 5] and experimental conditions [sprints 6 to 10]), and the difference in

203 the fastest sprint trial in each condition (i.e. fastest individual sprint at baseline minus fastest
204 individual sprint at experimental) were calculated.

205 Repeated measures ANOVA identified differences in sprint performance between each
206 condition, with treatment included as a between-subject factor. Greenhouse-Geisser epsilon
207 was reported where sphericity was violated, and post-hoc LSD tests were conducted where a
208 significant interaction was observed. Point-Biserial correlations (r_{pb}) were used to assess the
209 relationship between performance and categorical variables (i.e. sex, age, ability, sport
210 supplement use, frequency of sport supplement use, intention to use sport supplements, belief
211 manipulation scores). Data of the variables that correlated significantly with performances
212 were further analysed using repeated measures ANOVA and Multivariate ANOVA
213 (MANOVA). Given the possibility that differences between treatments may reflect the large
214 sample size and sampling variability (38), Cohen's d (d) effect sizes were calculated.
215 Differences between 0.2 and <0.5 were interpreted as a small effect, between 0.5 and <0.8 as
216 moderate, and ≥ 0.8 as large (13). Data are presented as mean \pm standard error of the mean
217 (SEM), with statistical significance accepted at $P \leq 0.05$.

218 **Results**

219 Participant demographics

220 No significant differences were observed between treatments for number of years training
221 ($F_{(2,573)} = 2.072$, $P = 0.127$), hours per week training ($F_{(2,580)} = 0.403$, $P = 0.669$), sex ($\chi^2 = 5.28$,
222 $P = 0.071$), supplement use ($\chi^2 = 2.32$, $P = 0.312$), frequency of supplement use ($\chi^2 = 6.50$, $P =$
223 0.370) and intention to use supplements ($\chi^2 = 4.65$, $P = 0.098$). Differences between treatments
224 were observed for age ($\chi^2 = 21.99$, $P = 0.001$), ability ($\chi^2 = 21.69$, $P = 0.001$) and sport played
225 ($\chi^2 = 225.76$, $P < 0.001$). Covariate analysis, adjusting for the differences in categorical

226 variables, revealed no effect on the outcome of the performance sprint data ($P > 0.05$). The
227 results of the subsequent analyses are therefore reported with unadjusted covariate data.

228 Reliability of sprint trials

229 Baseline sprints (i.e. trials 1 – 5) were associated with very high reliability in the positive-belief
230 treatment (ICC = 0.94), negative-belief treatment (ICC = 0.96) and control treatment (ICC =
231 0.90). Similar reliability coefficients were also observed for experimental sprints (i.e. trials 6 –
232 10) in the positive-belief treatment (ICC = 0.94), negative-belief treatment (TE = 0.94) and
233 control treatment (ICC = 0.94).

234 We also investigated the possibility that greater reliability was associated with fewer than 5
235 sprint trials. If for example, reliability between sprint trials 1 – 4 or 1 – 3 are more reliable than
236 1 – 5, this could reduce the error and improve the chances of finding a true effect of the
237 intervention on sprint performance. ICC's were however, similar for trials 1 – 4 (ICC range =
238 0.92 to 0.96) and 1 – 3 (ICC range = 0.93 to 0.96). Therefore, sprint trials 1 – 5 are reported in
239 the subsequent analysis.

240 Differences in baseline and experimental performance between treatments

241 No between-treatment differences were observed at baseline ($F_{(2,624)} = 0.149$, $P = 0.861$).
242 However, between-treatment differences were observed in experimental trials ($F_{(2,624)} = 5.879$,
243 $P = 0.001$). In the negative-belief treatment, performance was worse than at baseline ($-1.42 \pm$
244 0.15% , $P < 0.001$, $d = 0.56$), and also worse than performance in the positive-belief treatment
245 ($-1.04 \pm 0.28\%$, $P < 0.001$, $d = 0.34$) and in the control treatment ($-0.92 \pm 0.31\%$, $P < 0.001$, d
246 $= 0.32$). No differences were observed between the positive-belief and control treatments ($-$
247 $0.07 \pm 0.27\%$, $P = 0.696$, $d = 0.02$). Figure 1 illustrates the differences in performance for each
248 condition between treatments.

249 Correlations between performance and categorical variables

250 Point-Biserial correlations revealed a significant relationship between participant's intention to
251 use supplements and performance (average performance in each condition: $r_{pb} = 0.106$, $P =$
252 0.012 ; fastest performance difference between conditions: $r_{pb} = 0.101$, $P = 0.016$). No other
253 significant relationships were observed between other categorical variables for average
254 performance in each condition (sex $r_{pb} = -0.009$, $P = 0.819$; age $r_{pb} = 0.006$, $P = 0.891$; ability
255 $r_{pb} = -0.039$, $P = 0.353$; use of supplements $r_{pb} = 0.071$, $P = 0.078$; frequency of supplements
256 $r_{pb} = 0.075$, $P = 0.074$; belief manipulation scores $r_{pb} = -0.035$, $P = 0.563$) or fastest
257 performance between conditions (sex $r_{pb} = -0.014$, $P = 0.723$; age $r_{pb} = 0.005$, $P = 0.906$; ability
258 $r_{pb} = -0.042$, $P = 0.318$; use of supplements $r_{pb} = 0.075$, $P = 0.071$; frequency of supplements
259 $r_{pb} = -0.062$, $P = 0.135$; belief manipulation scores: $r_{pb} = 0.025$, $P = 0.677$; fastest performance:
260 $r_{pb} = 0.025$, $P = 0.677$).

261 Differences in baseline and experimental performance between supplement intention

262 Further analysis using repeated measures ANOVA identified differences in participant's repeat
263 sprint performance in each treatment by intention to use sport supplements (i.e. not intending;
264 $n = 174$; undecided; $n = 112$; and intending; $n = 284$). No differences between baseline and
265 experimental conditions were observed for participants in the positive-belief treatment
266 intending to use supplements ($0.28 \pm 0.14\%$, $P = 0.886$, $d = 0.01$). However, sprint performance
267 worsened for participants in the positive-belief treatment who were undecided about
268 supplement use ($-0.67 \pm 0.36\%$, $P = 0.039$; $d = 0.22$), and not intending to use sport
269 supplements ($-0.64\% \pm 0.25$, $P = 0.036$; $d = 0.23$; figure 2A). No differences in sprint
270 performance by intention to use supplements were observed in the negative-belief (figure 2B)
271 and control (figure 2C) treatments ($P > 0.05$).

272 Between-treatment differences in fastest performance by intention

273 Differences in fastest sprint performance and intention to use supplements were analysed using
274 MANOVA. The performance of participants intending to use supplements in the positive-belief
275 treatment was more positive compared to that of participants in the negative-belief treatment
276 ($1.29 \pm 0.37\%$, $P = 0.001$, $d = 0.51$) and control treatment ($0.90 \pm 0.41\%$, $P = 0.029$, $d = 0.33$).
277 Performance for participants not intending to use supplements in the negative-belief treatment
278 was worse compared than controls (negative-belief vs. controls = $-1.34 \pm 0.48\%$, $P = 0.005$, d
279 = 0.52). This trend was similar between the positive-belief and control treatment ($-0.91 \pm$
280 0.45% , $P = 0.060$; $d = 0.38$). No differences were observed for participant's undecided about
281 supplement use between all three treatments ($P > 0.05$; figure 3).

282 Within-treatment differences in fastest performance by intention

283 Differences in fastest sprint performance by intention to use supplements were observed in the
284 positive-belief treatment ($F_{(2,239)} = 4.952$, $P = 0.008$) but not in negative-belief treatment
285 ($F_{(2,197)} = 1.247$, $P = 0.290$) or control treatment ($F_{(2,131)} = 0.637$, $P = 0.530$). In the positive-
286 belief treatment, fastest sprint performance in experimental compared to baseline for
287 participants not intending to use supplements worsened by $-1.10\% \pm 0.30\%$, performance of
288 those undecided about supplement use worsened by $-0.64\% \pm 0.43\%$, while performance of
289 those intending to use supplements improved by $0.19\% \pm 0.24\%$ (figure 3). In the positive-
290 belief treatment, change in performance from baseline and experimental also differed
291 significantly between those participants intending to use supplements and those not intending
292 to use supplements ($1.29\% \pm 0.38\%$, $P = 0.003$, $d = 0.49$). No other within-treatment
293 differences in fastest sprint performance between baseline and experimental were observed
294 when classified by intention to use supplements ($P > 0.05$; figure 3).

295 **Discussion**

296 We aimed to replicate a previous study of placebo and nocebo effects in repeat sprint
297 performance (5), albeit with the inclusion of a no-treatment control and a larger sample. We
298 observed a mean nocebo effect in repeat sprint performance across the sample, but no mean
299 placebo effect when compared to a no-treatment control. This suggests that, while receiving a
300 purported harmful supplement significantly impaired performance, receiving a purported
301 beneficial supplement did not enhance it. This finding differs to those of Beedie et al. (5) who
302 reported significant placebo and nocebo effects in repeated sprinting.

303 Although no mean placebo effect was observed, data from the positive-belief treatment did
304 suggest that the performance of participants intending to use supplements improved to a greater
305 degree in the experimental conditions than the performance of participants not intending to use
306 supplements ($d = 0.49$, figure 3). These improvements were also greater than those observed
307 among participants of equivalent intention in the negative-belief treatment ($d = 0.51$) and
308 control treatment ($d = 0.33$). Given that effect sizes >0.2 are considered potentially beneficial
309 for sport performance (23), these improvements in repeat sprint performance are likely
310 meaningful for athletes. Furthermore, given that this relationship was observed only in the
311 positive-belief treatment is of particular importance, as it supports our hypothesis that intention
312 to use sports supplements might relate to placebo responding.

313 While intention to use supplements influenced the placebo response, this relationship was not
314 shown for prior supplement use ($r_{pb} = 0.071$, $P = 0.078$). We did however examine the effect
315 on performance of intention to use supplements and its interaction with prior supplement use.
316 Intention to use supplements was strongly associated with prior supplement use ($r_{pb} = 0.666$;
317 $P < 0.001$). This suggests that intention to use supplements is associated with prior supplement
318 use and may moderate an athlete's responsiveness to a placebo intervention. Although the
319 design of this research precluded a robust test of this relationship, it is an intriguing research
320 question that should be addressed in future research.

321 In consideration of the above, placebo responding is arguably a learned phenomenon. Research
322 has shown that placebo effects can be initiated via verbal instructions (creating an expectation
323 of a drug; (28)) and/or via repeat exposure to a drug with a subsequent placebo intervention
324 mirroring the action of that drug (9). Previous experiences of a drug are therefore remembered,
325 creating a memory of effective and ineffective treatments (29). This learning process is
326 manifest in specific brain regions, with expectations and conditioning cues mediating and
327 maintaining the turnover of, for example dopamine (19), and creating rewarding stimuli. On
328 this basis, for a placebo responsive athlete, a placebo induced improvement in performance is
329 the result of verbal information about the treatment (e.g. the suggestion that a supplement can
330 improve performance) and/or cued or contextual conditioning (e.g. repeated exposure to a real
331 treatment that results in treatment-like effects even when the treatment is replaced by a
332 placebo). The athlete then recalls previous experiences and information about the effectiveness
333 or ineffectiveness of the treatment, which shapes their subsequent intention to use it. This is
334 perhaps a reason why athletes intending to use supplements are more likely to use these
335 substances (17) and are arguably more likely to use other forms of performance enhancements
336 (26).

337 The finding that intention may influence the placebo effect has particular relevance to sports
338 practitioners aiming to improve an athlete's performance. Specifically, if improvements in
339 performance following administration of a treatment (e.g. caffeine, sodium bicarbonate, β -
340 alanine) are the result of both pharmacological and placebo effects (3), but the athlete does not
341 have a prior intention to use that treatment, it may not elicit a placebo response and the athlete
342 may not fully benefit from the treatment. Ultimately, a treatment may be more effective when
343 an athlete intends to use it than when they do not. Sport practitioners should therefore be aware
344 of an athlete's intentions towards a treatment prior to its administration, to ensure the

345 effectiveness of the treatment. This is also important in research, in which intentions towards
346 a treatment could likewise influence outcomes.

347 Any reference to the results of our study should take into account potential limitations. First,
348 we did not control for the presence of others or social support (e.g. cheering from teammates)
349 during the sprint trials, and this may have affected performance. Second, while participants
350 were asked to report on a Likert-type scale from 1 to 10 the degree to which they believed the
351 treatment influenced their performance, they were not specifically asked if they believed the
352 information they were given. We are therefore unable to assess the credibility of the belief-
353 manipulation. Finally, the use of self-reported sport supplement use may not be reliable, as
354 there may be differences between what athletes' report and what they actually think and/or do.

355 Given that previous studies have used expensive and complex techniques such as positron
356 emission tomography (2) and genotyping (18) to identify placebo responders/non-responders,
357 a self-report measure could provide a cost-effective and practical alternative. Future research
358 should aim to further explore the impact of intention on the effects of legitimate sports
359 supplements, and how this could influence an athlete's decision to use other forms of
360 performance enhancements (e.g. doping). This understanding could enhance treatments, and
361 inform athlete education and anti-doping strategy (26).

362 **Acknowledgements**

363 We would like to thank participants for their time and effort. We also greatly appreciate the
364 constructive comments by the anonymous reviewers, which improved the quality of the
365 manuscript. Authors received no external funding for this research and declare no conflicts of
366 interest. The results of the present study do not constitute endorsement by ACSM. The results
367 of this study are presented clearly, honestly and without fabrication, falsification or
368 inappropriate data manipulation

369 **References**

- 370 1. American Psychological Association. Ethical principles of psychologists and code of
371 conduct. *Amer Psychol.* 2010;57(12):1060-73.
- 372 2. Atlas LY, Wager TD. A meta-analysis of brain mechanisms of placebo analgesia:
373 consistent findings and unanswered questions. *Handb Exp Pharmacol.* 2014;225:37-
374 69.
- 375 3. Beedie CJ. Placebo effects in competitive sport: qualitative data. *J Sports Sci Med.*
376 2007;6(1):21-8.
- 377 4. Beedie CJ. All in the mind? Pain, placebo effect, and ergogenic effect of caffeine in
378 sports performance. *Open Access J Sports Med.* 2010;1:87-94.
- 379 5. Beedie CJ, Coleman DA, Foad AJ. Positive and negative placebo effects resulting from
380 the deceptive administration of an ergogenic aid. *Int J Sport Nutr Exerc Metab.*
381 2007;17(3):259-69.
- 382 6. Beedie CJ, Foad AJ. The placebo effect in sports performance: a brief review. *Sports*
383 *Med.* 2009;39(4):313-29.
- 384 7. Beedie CJ, Foad AJ, Coleman DA. Identification of placebo responsive participants in
385 40km laboratory cycling performance. *J Sports Sci Med.* 2008;7(1):166-75.
- 386 8. Beedie CJ, Stuart EM, Coleman DA, Foad AJ. Placebo effects of caffeine on cycling
387 performance. *Med Sci Sports Exerc.* 2006;38(12):2159-64.
- 388 9. Benedetti F, Amanzio M, Rosato R, Blanchard C. Nonopioid placebo analgesia is
389 mediated by CB1 cannabinoid receptors. *Nat Med.* 2011;17(10):1228-30.
- 390 10. Carlino E, Benedetti F. Different contexts, different pains, different experiences.
391 *Neuroscience.* 2016;338:19-26.
- 392 11. Carlino E, Piedimonte A, Benedetti F. Nature of the placebo and nocebo effect in
393 relation to functional neurologic disorders. *Handb Clin Neurol.* 2017;139:597-606.

- 394 12. Clark VR, Hopkins WG, Hawley JA, Burke LM. Placebo effect of carbohydrate
395 feedings during a 40-km cycling time trial. *Med Sci Sports Exerc.* 2000;32(9):1642-7.
- 396 13. Cohen J. A power primer. *Psychol Bull.* 1992;112(1):155.
- 397 14. Cross MR, Brughelli M, Brown SR et al. Mechanical properties of sprinting in elite
398 Rugby union and Rugby league. *Int J Sports Physiol Perform.* 2015;10(6):695-702.
- 399 15. Ferreira TN, Sabino-Carvalho JL, Lopes TR et al. Ischemic preconditioning and
400 repeated sprint swimming: A placebo and nocebo study. *Med Sci Sports Exerc.*
401 2016;48(10):1967-75.
- 402 16. Foad AJ, Beedie CJ, Coleman DA. Pharmacological and psychological effects of
403 caffeine ingestion in 40-km cycling performance. *Med Sci Sports Exerc.*
404 2008;40(1):158-65.
- 405 17. Goulet C, Valois P, Buist A, Cote M. Predictors of the use of performance-enhancing
406 substances by young athletes. *Clin J Sport Med.* 2010;20(4):243-8.
- 407 18. Hall KT, Lembo AJ, Kirsch I et al. Catechol-O-methyltransferase val158met
408 polymorphism predicts placebo effect in irritable bowel syndrome. *PLoS One.*
409 2012;7(10):e48135.
- 410 19. Hall KT, Loscalzo J, Kaptchuk TJ. Genetics and the placebo effect: the placebo.
411 *Trends Mol Med.* 2015;21(5):285-94.
- 412 20. Haugen T, Buchheit M. Sprint running performance monitoring: Methodological and
413 practical considerations. *Sports Med.* 2016;46(5):641-56.
- 414 21. Hopker JG, Foad AJ, Beedie CJ, Coleman DA, Leach G. Placebo effect of an inert gel
415 on experimentally induced leg muscle pain. *Open access journal of sports medicine.*
416 2010;1:215-21.
- 417 22. Hopkins WG. Spreadsheets for Analysis of Validity and Reliability. *Sportsci.*
418 2015;19:26-42.

- 419 23. Hopkins WG. Estimating sample size for magnitude-based inferences. *Sportsci.*
420 2016;10:63-9.
- 421 24. Hopkins WG, Hawley JA, Burke LM. Design and analysis of research on sport
422 performance enhancement. *Med Sci Sports Exerc.* 1999;31(3):472-85.
- 423 25. Hrobjartsson A, Kaptchuk TJ, Miller FG. Placebo effect studies are susceptible to
424 response bias and to other types of biases. *J Clin Epidemiol.* 2011;64(11):1223-9.
- 425 26. Hurst P, Foad AJ, Coleman DA, Beedie CJ. Development and validation of the Sports
426 Supplements Beliefs Scale Perform Enhanc Health. In press.
- 427 27. Judd CM, McClelland GH, Ryan CS. *Data analysis: A model comparison approach.*
428 Routledge; 2011, 295 p.
- 429 28. Kam-Hansen S, Jakubowski M, Kelley JM et al. Altered placebo and drug labeling
430 changes the outcome of episodic migraine attacks. *Sci Transl Med.*
431 2014;6(218):218ra5.
- 432 29. Klinger R, Colloca L, Bingel U, Flor H. Placebo analgesia: clinical applications. *Pain.*
433 2014;155(6):1055-8.
- 434 30. Leys C, Ley C, Klein O, Bernard P, Licata L. Detecting outliers: Do not use standard
435 deviation around the mean, use absolute deviation around the median. *Journal of*
436 *Experimental Social Psychology.* 2013;49(4):764-6.
- 437 31. Maganaris C, Collins D, M S. Expectancy Effects and Strength Training: Do Steroids
438 Make a Difference? *The Sport Psychologist.* 2000;14:272–78.
- 439 32. McClung M, Collins D. “Because I know it will!”: placebo effects of an ergogenic aid
440 on athletic performance. *Journal of Sport and Exercise Psychology.* 2007;29(3):382-
441 94.

- 442 33. Ntoumanis N, Ng JY, Barkoukis V, Backhouse S. Personal and psychosocial predictors
443 of doping use in physical activity settings: a meta-analysis. *Sports Med.*
444 2014;44(11):1603-24.
- 445 34. Ross R, Gray CM, Gill JM. Effects of an injected placebo on endurance running
446 performance. *Med Sci Sports Exerc.* 2015;47(8):1672-81.
- 447 35. Saunders B, de Oliveira LF, da Silva RP et al. Placebo in sports nutrition: a proof-of-
448 principle study involving caffeine supplementation. *Scand J Med Sci Sports.* In press.
- 449 36. Schimpchen J, Skorski S, Nopp S, Meyer T. Are "classical" tests of repeated-sprint
450 ability in football externally valid? A new approach to determine in-game sprinting
451 behaviour in elite football players. *J Sports Sci.* 2016;34(6):519-26.
- 452 37. Spencer M, Bishop D, Dawson B, Goodman C. Physiological and metabolic responses
453 of repeated-sprint activities: specific to field-based team sports. *Sports Med.*
454 2005;35(12):1025-44.
- 455 38. Sullivan GM, Feinn R. Using effect size-or why the P value is not enough. *J Grad Med*
456 *Ed.* 2012;4(3):279-82.
- 457 39. Trojian TH, Beedie CJ. Placebo effect and athletes. *Curr Sports Med Rep.*
458 2008;7(4):214-7.
- 459

Table 1. Demographics of participants between treatments

		Positive	Negative	Control	Overall
	n =	288	232	192	712
Gender (%)	Male	83.1	76.9	71.0	78.0
	Female	16.9	23.1	29.0	22.0
Age (%)	18 to 24	66.7	65.0	79.0	69.4
	25 to 34	29.6	30.0	18.8	26.8
	35 to 44	3.7	5.1	2.3	3.8
Sport (%)	Rugby Union	46.2	42.7	22.3	39.0
	Soccer	42.9	36.9	44.1	41.3
	Field Hockey	5.3	8.9	2.8	5.8
	Other	5.6	11.6	30.7	13.9
Ability (%)	Club	25.5	35.4	21.1	27.5
	County	39.9	38.8	30.4	37.0
	Regional	25.9	19.6	32.7	25.7
	National	8.7	6.2	15.8	9.8
Intention to use sport supplements (%)	Not intending	23.9	33.5	35.6	30.0
	Undecided	21.6	18.9	18.1	19.8
	Intending	54.5	47.6	46.3	50.2
Use of Supplements (%)	Yes	51.1	50.9	52.7	51.5
	No	48.9	49.1	47.4	48.5
Frequency of supplement use (%)	Daily	24.1	26.6	26.2	25.5
	Weekly	22.6	21.0	24.4	22.5
	Monthly	4.4	3.3	1.8	3.4
	Never	48.9	49.1	47.6	48.6
Mean ± SEM	Years training	10.77 ± 0.38	10.94 ± 0.59	9.68 ± 0.45	10.68 ± 0.24
	Hours per week training	6.13 ± 0.25	5.93 ± 0.25	5.84 ± 0.30	5.9 ± 0.15
	Amount of supplements used	1.14 ± 0.10	1.11 ± 0.10	1.20 ± 0.13	1.09 ± 0.06

SEM, standard error of the mean

461 **Figure captions**

462 **Figure 1.** Average performance in each condition between treatments. Note: *baseline vs.
463 experimental for negative-belief = $P < 0.05$; **positive-belief and control vs. negative-belief =
464 $P < 0.05$.

465

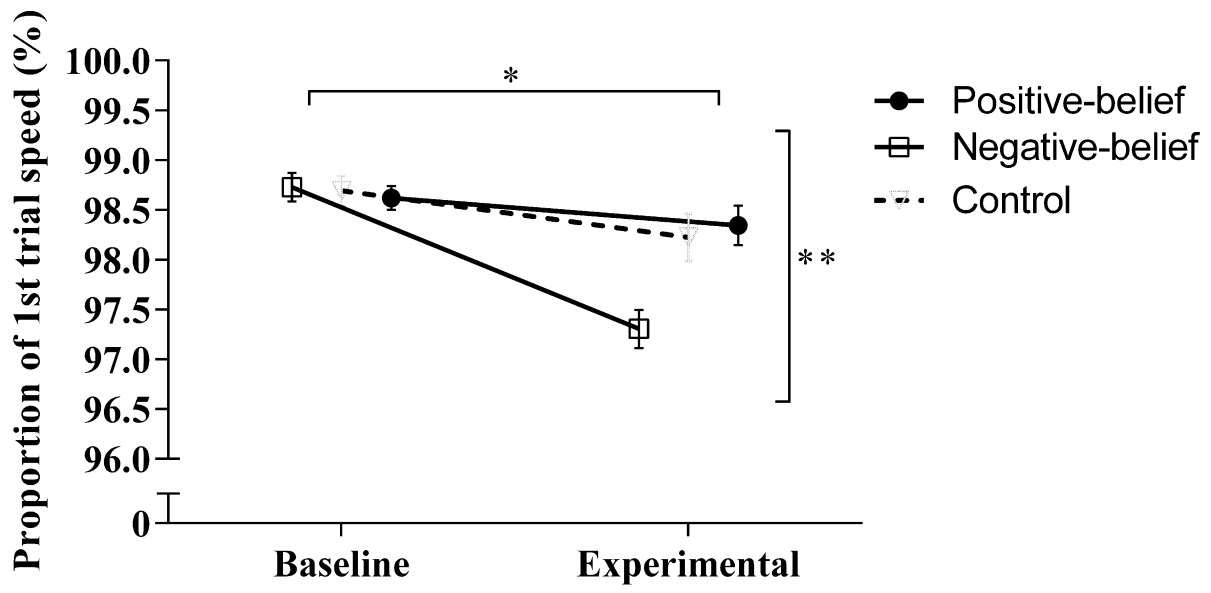
466 **Figure 2.** Average performance in condition by each treatment separated by participants'
467 intention to use sport supplements in the next three months. **A.** Positive-belief treatment. Note:
468 *Baseline vs. Experimental for those not intending to use supplements = $P < 0.05$; **intending
469 to use supplements vs. not intending to use supplements = $P < 0.05$. **B.** Negative-belief
470 treatment. Note: *baseline vs. experimental for those not intending, undecided and intending
471 to use supplements = $P < 0.05$. **C.** No-treatment control.

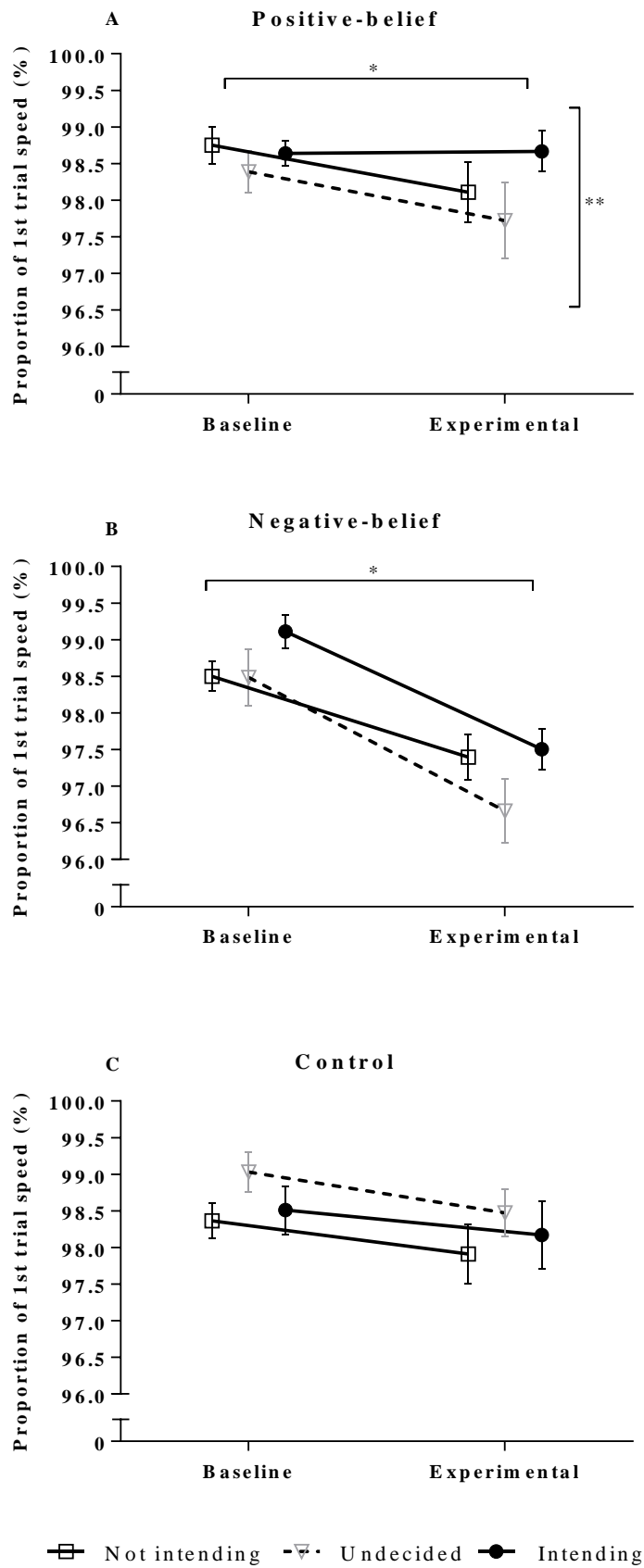
472

473 **Figure 3.** Differences in fastest performance between conditions, grouped by intention to use
474 sport supplements. Note: *control vs. positive-belief and negative-belief = $P < 0.05$, **positive-
475 belief vs. negative-belief = $P < 0.05$, †positive-belief intention vs. positive-belief no intention
476 = $P < 0.05$

477

478 **Figure 1**





480 **Figure 3.**

