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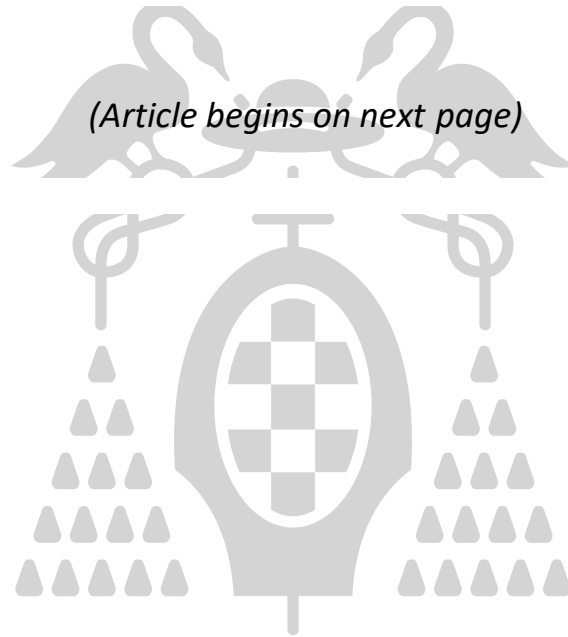


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2

3 **Effect of acute sodium bicarbonate and caffeine co-ingestion on repeated sprint**
4 **performance in recreationally trained individuals: a randomized controlled trial**

5

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34 **ABSTRACT**

35 **Introduction.** The acute and isolated ingestion of sodium bicarbonate and caffeine improves
36 performance and delays fatigue in high-intensity tasks. However, it remains to be elucidated if
37 the co-ingestion of both dietary supplements stimulates a summative ergogenic effect. This
38 study aimed to examine the effect of the acute co-ingestion of sodium bicarbonate and caffeine
39 on repeated sprint performance.

40 **Methods.** Twenty-five trained participants (age: 23.3 ± 4.0 years; sex(female/male): 12/13; body
41 mass: 69.6 ± 12.5 kg) participated in a randomized, double-blind, placebo-controlled and cross-
42 over study. Participants were assigned to four conditions: a) sodium bicarbonate and caffeine
43 ($\text{NaHCO}_3 + \text{CAF}$); b) sodium bicarbonate (NaHCO_3); c) caffeine (CAF); d) placebo (PLA).
44 Thus, they ingested 0.3 g/kg of NaHCO_3 , 3 mg/kg of caffeine or placebo. Then, participants
45 performed 4 Wingate tests (Wt), consisting of a 30-s all-out sprint against an individualized
46 resisted load, interspersed by a 1.5 min rest period among sprints.

47 **Results.** Peak (W_{peak}) and mean (W_{mean}) power output revealed a supplement and sprint
48 interaction effect ($P=0.009$ and $P=0.049$, respectively). Compared to placebo, $\text{NaHCO}_3 + \text{CAF}$
49 and NaHCO_3 increased W_{peak} performance in Wt3 (3%, $P=0.021$) and Wt4 (4.5%, $P=0.047$),
50 while NaHCO_3 supplementation increased W_{mean} performance in Wt3 (4.2%, $P=0.001$). In Wt1
51 CAF increased W_{peak} (3.2%, $P=0.054$) and reduced time to W_{peak} (-8.5%; $P=0.008$). Plasma
52 lactate showed a supplement plus sprint interaction ($P<0.001$) when NaHCO_3 was compared to
53 caffeine (13%, $P=0.031$) and placebo (23%, $P=0.021$).

54 **Conclusions.** To summarize, although the isolated ingestion of caffeine and sodium bicarbonate
55 improved repeated sprint performance, the co-ingestion of both supplements did not stimulate a
56 synergic ergogenic effect.

57

58 **Keywords:** sports performance, sports nutrition, ergogenic substances, caffeine, sodium
59 bicarbonate, sprint.

60 INTRODUCTION

61 Sodium bicarbonate (NaHCO_3) and caffeine supplementation are widespread among athletes of
62 different sports modalities since both supplements improve high-intensity tasks that last
63 between 30 s to 8 min^{1,2}. Since NaHCO_3 and caffeine could enhance performance through
64 different physiological mechanisms²⁻⁴, the co-ingestion of both supplements could stimulate an
65 additive ergogenic effect. However, the interaction between NaHCO_3 and caffeine has been
66 scarcely studied in repeated sprint performance⁵ despite being a critical performance component
67 in several sports.

68 Caffeine is one of the most commonly used ergogenic aid whose effect mainly occurs in
69 the CNS by antagonizing adenosine receptors reducing fatigue and perceived effort, increasing
70 alertness and vigour, and facilitating muscle fibers recruitment during muscle contraction².
71 Besides, caffeine may promote intracellular calcium ion (Ca^{2+}) mobilization, stimulating force
72 production⁶ and delaying fatigue caused by a gradual reduction in Ca^{2+} bioavailability⁷.
73 NaHCO_3 ergogenicity also occurs in skeletal muscle where increased extracellular buffering by
74 stimulating higher extracellular pH and base excess, leading to an increase of H^+ and lactate co-
75 transport out of the exercising muscle cells⁸. Thus, the alkalizing influence of NaHCO_3 in blood
76 and muscle during exercise, increasing glycolytic rates⁸ and higher rates of ATP re-synthesis
77 and Ca^{2+} utilization to sustain exercise demands delaying the muscular loss of ability to
78 generate force and power^{3,4}. Although both supplements may delay fatigue effects, particularly
79 important in repeated effort, the combined effect of both supplements have been scarcely
80 explored.

81 Considering the isolated effect of caffeine and NaHCO_3 on performance and that these
82 supplements seem to promote their ergogenicity through different mechanisms of action, it can
83 be argued that the co-ingestion of both supplements could promote a synergic effect. This idea
84 was explored in Felipe et al.⁹ study where the co-ingestion of caffeine and NaHCO_3 promoted
85 additive effects in ten male judokas who performed Special Judo Fitness Tests after ingesting 6
86 mg/kg of caffeine 50 min before exercise and 0.3 g/kg of NaHCO_3 divided in three doses

87 ingested at 120, 90 and 60 minutes before⁹. However, this effect was not observed using doses
88 of 3-6 mg/kg of caffeine and 0.3 g/kg of NaHCO₃ provided in one or several doses from 120 to
89 60 min before the trial in eight trained rowers after a 2000 m rowing task¹⁰, in twelve elite
90 rowers after a 6 min rowing test¹¹, in ten trained cyclist after 3-km time trial¹², in thirteen non-
91 cycling trained individuals after cycling to volitional exhaustion¹³, in six elite swimmers after a
92 2x200m swimming test¹⁴ or in eight karate athletes after a karate-specific aerobic test¹⁵.

93 Nevertheless, it should be considered that NaHCO₃ supplementation protocols in some of
94 the studies^{13,14} resulted in high incidence and severity of side effects (e.g., gastrointestinal
95 discomfort) that undoubtedly affected performance. Moreover, the sample sizes of these studies
96 were small, ranging from 6 to 13 participants in most studies. Additionally, since caffeine may
97 increase mood¹⁶, motivation¹⁷ and reduce fatigue¹⁸, it can be argued that the inter-individual
98 response to caffeine may be attributable to variances in these variables¹⁹. Therefore, further
99 work is required to determine whether co-supplementation of caffeine and NaHCO₃ produces
100 larger ergogenic effects than isolated supplementation particularly in repeated high-intensity
101 efforts where evidence is lacking since caffeine may cause an ergogenic effect during the first
102 sprint² and sodium bicarbonate after at least two of them²⁰. Thus, this study aimed to examine
103 the effect of the acute co-ingestion of NaHCO₃ and caffeine on repeated sprint performance in
104 both male and female participants.

105

106 **MATERIALS AND METHODS**

107 **Participants**

108 Twenty-five individuals (age: 23.3±4.0 years; sex (female/male): 12/13; body mass: 69.6±12.5
109 kg) participated in this study. All participants were recreationally trained (training experience:
110 3.1±1.2 years and 4.5±1.2 days/week) habituated to anaerobic exercise and to caffeine
111 consumption (81±70 mg/day). Besides, participants had no diagnosis of musculoskeletal,
112 neurological, immunological or cardio-metabolic disorders, a training experience of at least 8
113 months before the intervention, performing 3 days per week during the previous 3 months, and

114 did not take any medication, drug, stimulant or any other sports supplement during the trial. Six
115 of twelve female participants initiated the trial during the follicular phase of their menstrual
116 cycle.

117 Procedures, potential risks or discomfort associated with the experiments were explained
118 to participants, who then gave their written informed consent. The Ethics Committee of
119 Investigation and Animal Experimentation from the University of Alcalá approved the
120 experiment (CEIP/HU/2021/1/006), which is in accordance with the Declaration of Helsinki.

121

122 **Experimental design**

123 The study design was randomized, double-blind, cross-over and placebo-controlled. Each
124 participant reported five times to the laboratory (Faculty of Medicine and Health Sciences,
125 Laboratory: 044.01.047.0). Participants underwent preliminary questionnaires of dietary and
126 physical activity habits and body composition assessment during visit one, and a familiarization
127 session where they experienced all tests performed in the trials.

128 During visits two to five, volunteers reported to the laboratory at the same time of day
129 (± 30 min). They participated in four trials separated by at least 72h to allow a complete
130 recovery and washout period. Participants were assigned to four conditions: a) Sodium
131 bicarbonate and caffeine ($\text{NaHCO}_3 + \text{CAF}$); b) sodium bicarbonate (NaHCO_3); c) caffeine
132 (CAF); d) placebo (PLA). The order of the trials was randomized in sequence according to the
133 experimental condition for each participant (www.randomized.org). An external researcher was
134 responsible for elaborating the alphanumeric code assigned to each sequence to blind
135 participants and researchers during the trials. The codes were unveiled after statistical analysis.

136

137 **Experimental protocol**

138 ***Body composition, dietary and physical activity habits***

139 As previously reported elsewhere, body composition was assessed using electric bioimpedance
140 (Tanita BC-418, Tanita Corporation of America Inc. IL, USA). Dietary habits were analyzed

141 using a 24h dietary recall and the software MyFitnessPal and DIAL (Alce Ingeniería, Madrid,
142 Spain), while physical activity habits were evaluated using the International Physical Activity
143 Questionnaire (IPAQ). Twenty-four hours before the familiarization session and until the end of
144 the trial, participants were encouraged to refrain from caffeine, stimulants and alcohol intake.
145 Besides, 24h before each visit (familiarization and trials), participants refrained from strenuous
146 exercise and were asked to follow a similar sleep and dietary pattern.

147

148 ***Supplementation protocol***

149 The supplementation protocol started 120 min before the trial. Participants ingested NaHCO₃
150 (0.3 g/kg of body mass) or placebo (3 mg/kg, of maltodextrin, HSN, Granada, Spain) on two
151 occasions at 120 and 90 min before the trial, consuming 0.15 g/kg of NaHCO₃ or 1.5 mg/kg of
152 maltodextrin on each time. This supplementation strategy was designed to minimize
153 gastrointestinal problems⁹. Then, 60 minutes before the trial, participants ingested caffeine (3
154 mg/kg, HSN, Granada, Spain) or placebo (3 mg/kg, maltodextrin).

155 Supplements were dissolved in 150 ml of tap water and a flavoring with no calories was added
156 to mask the supplements' flavor and smell (MyProtein, Northwich, UK). The beverages were
157 provided in opaque shaker bottles.

158

159 ***Repeated sprint test (4 x 30s) and plasma lactate***

160 Participants performed a standardized warm-up of 5 min of pedaling in an isoinertial cycle
161 ergometer (Monark LC6, Monark. Vansbro, Suecia) at 70 W without any resisted load. Then,
162 participants performed four Wingate tests (Wt), interspersed by a passive 1.5 min rest period
163 among Wt, each consisting of a 30-s all-out sprint against an individualized resisted load (0.075
164 Nm/kg of body mass). Participants were given verbal encouragement throughout the tests. Peak
165 and mean power, time to reach peak power and fatigue index (FI) were obtained. FI was
166 calculated by taking the minimum power away from the peak power and then dividing it by the
167 peak power and multiplying it by 100.

168 The number of Wingate tests performed to explore the potential interaction between
169 caffeine and sodium bicarbonate was selected based on the potentially expected ergogenic
170 effect of caffeine during the first sprint^{2,21} and NaHCO₃ during the third and fourth sprints²⁰.
171 Moreover, the passive rest interval between sprints was selected to stimulate a pronounced
172 decrease in pH levels to emphasize the potential buffering capacity of NaHCO₃²².

173 Before and immediately after Wt1 and immediately after Wt4, capillary blood samples
174 were collected for the assessment of plasma lactate (mmol/l) (Lactate Pro2, ProTM 2 LT-1710
175 Instrument, Arkray Fatory Inc., KDK Corporation, Shiga, Japan).

176

177 *Questionnaires and scales*

178 As previously reported, participants' mood was assessed using a reduced version of the profile
179 of mood states questionnaire (POMS) and subjective vitality scale (SVS)²¹. Participants graded
180 a set of 29 items related to the mood on a Likert scale from 0 (not at all) to 4 (extremely) in
181 reply to the question "How do you feel at this moment?" to assess six scales: tension,
182 depression, anger, vigor, fatigue and confusion. Moreover, participants' vitality was evaluated
183 using the subjective vitality scale²³. Using a 7-point Likert scale where 1 means "total
184 disagreement" and 7 means "total agreement", participants reported their subjective feelings of
185 energy and vitality.

186 At the end of the trials, participants had to complete a side effects questionnaire about
187 their perception of power, endurance, energy and exertion, heart, muscular and gastrointestinal
188 discomfort²⁴. Additionally, a specific question to evaluate the blinding procedure was also
189 included.

190

191 **Statistical Analysis**

192 The sample size calculation revealed that 20 participants were sufficient for the purpose of the
193 study to show an effect size of 0.35 ($\alpha=0.05$; $1-\beta=0.80$) (v3.1, G*power, Dusseldorf University,
194 Germany); finally, 25 participants were recruited.

195 Data collected in the study were analyzed using the statistical package SPSS v27.0 (SPSS
196 Inc., Chicago, IL, USA) and figures were generated using GraphPad Prism (v8, GraphPad
197 Software Inc., La Jolla, CA, USA). Initially, statistical analysis was conducted comparing sex
198 (male vs female), however, due to the lack of difference found, both sex groups were condensed
199 into one and treated as a single group. Firstly, Shapiro-Wilks was used to test the normality of
200 the data ($P > 0.05$). Plasma lactate was assessed using a two-way ANOVA for repeated measures
201 according to supplement (NaHCO₃+CAF, NaHCO₃, CAF and PLA) and time (Baseline, Wt1
202 and Wt4). Holm-Bonferroni correction was used as a *post hoc* test when significant differences
203 were detected.

204 Furthermore, body composition, dietary and physical habits were assessed using one-way
205 ANOVA. The Q the Cochran test was used to detect differences in blinding supplement success
206 before and after each trial and the side effects caused by both sports supplements

207 Values are reported as mean \pm standard deviation (SD). The significance level was set at
208 $P \leq 0.05$. Effect size (ES) was calculated as partial eta squared statistic (η_p^2) for the two-way
209 repeated measures and Hedges's (g) for partial comparisons based on the following criteria:
210 trivial (0–0.19), small (0.20–0.49), medium (0.50–0.79) and large (0.80 and greater) (Cohen,
211 1992).

212

213 **RESULTS**

214 No statistically significant differences among experimental conditions were found regarding
215 body composition, dietary or physical activity habits (Table 1). Differences in body
216 composition, dietary and physical activity habits on each experimental group according to sex is
217 shown in Supplementary Table 1.

218

219 *Repeated sprint test (4 x 30s) and lactate*

220 Differences in peak and mean power output, time to reach peak power output and fatigue index
221 are shown in Figure 1. Differences in repeated sprint performance after the four
222 supplementation protocols according to sex is shown in Supplementary Figure 1.

223 Peak power output (W_{peak}) revealed a supplement ($P=0.029$, $\eta_p^2=0.133$) and supplement
224 plus sprint interaction effect ($P=0.009$, $\eta_p^2=0.107$). $\text{NaHCO}_3+\text{CAF}$ supplementation increased
225 W_{peak} in Wt3 (3.0%, $P=0.021$, $g=0.182$) and Wt4 (4.5%, $P=0.047$, $g=0.303$) compared to
226 placebo. Similarly, compared to placebo, NaHCO_3 supplementation increased W_{peak} in Wt3
227 (3.7%, $P=0.032$, $g=0.178$) and Wt4 (6.8%, $P=0.042$, $g=0.298$). While CAF supplementation
228 showed a W_{peak} increase in Wt1 compared to placebo (3.2%, $P=0.054$).

229 Mean power output (W_{mean}) revealed a supplement ($P=0.040$, $\eta_p^2=0.201$) and supplement
230 plus sprint interaction effect ($P=0.049$, $\eta_p^2=0.101$). NaHCO_3 supplementation increased W_{mean}
231 in Wt3 (4.2%, $P=0.001$, $g=0.184$). No other partial difference was detected.

232 Time to W_{peak} did not report statistically significant supplement or supplement plus sprint
233 effects (figure 1). However, in Wt1, $\text{NaHCO}_3+\text{CAF}$ (-10.3%; $P=0.015$, $g=0.475$), NaHCO_3 (-
234 6.7%; $P=0.045$, $g=0.311$) and CAF (-8.5%; $P=0.008$, $g=0.271$) supplementation reduced time to
235 W_{peak} compared to placebo. Also, in Wt3, NaHCO_3 supplementation reduced time to W_{peak}
236 compared to placebo (-7.3%; $P=0.045$, $g=0.414$).

237 Fatigue Index did not report statistically significant supplement or supplement plus sprint
238 effects (figure 1). However, in Wt4, FI increased in $\text{NaHCO}_3+\text{CAF}$ (5.4%, $P=0.050$, $g=0.245$),
239 NaHCO_3 (9.3%, $P=0.037$, $g=0.421$) and CAF supplementation (7.7%, $P=0.049$, $g=0.328$)
240 compared to placebo.

241 Finally, lactate showed a supplement plus sprint interaction ($P<0.001$, $\eta_p^2=0.244$, Figure
242 1). Particularly, differences were found immediately after the Wt4, when comparing
243 $\text{NaHCO}_3+\text{CAF}$ to caffeine (17%, $P=0.002$, $g=0.630$) and placebo (28%, $P=0.004$, $g=0.590$).
244 These differences were also found when NaHCO_3 was compared to caffeine (13%, $P=0.031$,
245 $g=0.547$) and placebo (23%, $P=0.021$, $g=0.90$).

246

247 *Questionnaires and scales*

248 No statistical differences were found in depression, anger, vigor, fatigue, confusion or SVS
249 (Table 2). However, in tension, CAF showed an increase compared to the rest of the
250 supplementation protocols ($P=0.020$, $\eta_p^2=0.155$).

251 The side effects questionnaire revealed an interaction effect for gastrointestinal ($P=0.003$,
252 $\eta_p^2=0.262$) and muscular discomfort ($P=0.005$, $\eta_p^2=0.256$). Partial comparison revealed that,
253 immediately after the trials, $\text{NaHCO}_3+\text{CAF}$ increased gastrointestinal discomfort compared to
254 CAF (45%; $P=0.005$, $g=0.910$) and placebo (57%; $P=0.021$, $g=1.29$), while NaHCO_3 increase
255 discomfort compared to CAF (42%; $P=0.039$, $g=0.869$) and placebo (54%; $P=0.012$, $g=1.292$).
256 Moreover, CAF increased muscular discomfort compared to $\text{NaHCO}_3+\text{CAF}$ (21%; $P=0.044$,
257 $g=0.329$), NaHCO_3 (48%; $P=0.021$, $g=0.788$) and placebo (32%; $P=0.041$, $g=0.263$). Also,
258 higher energy perception was found in $\text{NaHCO}_3+\text{CAF}$ (23%; $P=0.012$, $g=0.525$) and CAF
259 (15%; $P=0.047$, $g=0.317$) compared to the placebo.

260 Finally, 72% (18 of 25) of participants correctly guessed when they ingested bicarbonate,
261 while 60% (15 of 25) correctly guessed when they ingested caffeine.

262

263 **DISCUSSION**

264 The purpose of this study was to examine the effect of the acute co-ingestion of NaHCO_3 and
265 caffeine on repeated sprint performance. Our results suggest that the isolated ingestion of
266 NaHCO_3 increased peak and mean power output, fatigue index and reduced time to reach peak
267 power in the third and fourth Wingate tests, while caffeine reduced time to reach peak power
268 only in the first sprint. Despite the ergogenic effects promoted by isolated ingestion, the co-
269 ingestion of caffeine and NaHCO_3 did not provide summative effects in repeated sprint
270 performance.

271

272 Power output (W_{peak} and W_{mean}) is commonly measured during a single or repeated sprint
273 task using Wingate tests. Our study shows that NaHCO_3 ingestion, isolated or combined with

274 caffeine, increases W_{peak} to a similar extent in the third and fourth Wingate test compared to
275 placebo. In a previous meta-analysis, Grgic²⁰ showed that NaHCO_3 intake increases W_{peak} only
276 in a third of fourth performed Wingate test. However, this evidence was supported only by two
277 studies consisting of four Wingate tests in the upper-body interposed by 6 min of rest in judo
278 athletes²⁵ and three lower-body Wingate tests interposed by 6 min of rest in recreationally
279 trained males²⁶. In contrast, other studies do not support this idea²⁷⁻²⁹. In previous studies
280 developed by Zabala et al.^{27,28}, NaHCO_3 did not improve W_{peak} in 9-10 elite male bicycle
281 motocross riders who performed three Wingate tests with 15-30 min of rest among them.
282 Similarly, Zinner et al.²⁹ did not find statistical differences in W_{peak} in 11 aerobically well-
283 trained men after four Wingates tests performed in the lower-body with 5 min of rest. In our
284 study, the recovery time among sprints was considerably lower compared to previous studies,
285 1.5 minutes among Wingate tests. This shorter recovery time may have elicited a higher muscle
286 acidosis which seems critical to facilitate the NaHCO_3 effect on the attenuation of W_{peak}
287 diminution through repeated sprints.

288 In our study, NaHCO_3 intake increased W_{mean} in the third Wingate test, an effect that was
289 reported when this supplement was ingested alone. Besides, in the fourth Wingate test,
290 NaHCO_3 promoted a non-statistically significant increase in W_{mean} by 2-3%. Previous studies
291 have reported an increase in W_{mean} in the third and fourth Wingate tests performed in the upper-
292 body²⁵ and lower-body²⁹. However, other studies did not find these differences²⁶⁻²⁸. The
293 comparison of these studies revealed that those studies with a short rest interval among sprints
294 (3-5 min) observed a significant effect of NaHCO_3 on W_{mean} ^{25,29}, while the studies in which a
295 more prolonged rest period was allowed among sprints (6-30 min) did not found ergogenic
296 effect on this performance parameter. Thus, as it occurs with W_{peak} , these studies together with
297 our results, support the idea that a short rest period (<5 min) is needed to find an ergogenic
298 effect of NaHCO_3 on W_{mean} during repeated sprint tests.

299 Peak power output represents the ability to produce mechanical power in the shorter
300 period possible, whereas W_{mean} reflects the average power produced during 30 seconds. W_{mean}

301 could be seen as a reflection of the active muscle endurance, and although a more pronounced
302 ergogenic effect of NaHCO₃ could be expected on muscle endurance (W_{mean}) compared to
303 W_{peak} ³⁰, a similar increase in performance of W_{peak} and W_{mean} was found in our study. This may
304 be explained due to the fact that the repeated sprint protocol in which four Wingate tests were
305 performed interposed by 1.5 min of rest among sprints caused a considerable amount of fatigue
306 among participants. This is supported by the increase in FI and plasma lactate observed at the
307 end of the repeated sprint protocol. We observed that the three supplement conditions increased
308 FI by 5-9 % in Wt4. FI is dependent on the peak and minimum power produced in a sprint,
309 since acute NaHCO₃ intake showed to preserve W_{peak} in the Wt4, it was not surprising to find
310 an elevated FI in this experimental condition and time. Nonetheless, our results are not in line
311 with other previous studies^{28,31}, maybe due to differences in the number of sprints performed (3
312 vs 4), the resting protocol used (15-30 min vs 1.5 min) and the sample size compared to our
313 study. On the other hand, caffeine supplementation does not affect W_{peak} in the Wt4 but
314 promotes an improvement in FI in this sprint. Hence, the ergogenic effect of caffeine on FI may
315 be explained by enlarging the range from the peak and minimum power produced in this sprint.

316 Furthermore, plasma lactate showed a supplement plus sprint interaction observed after
317 NaHCO₃ supplementation after the fourth sprint. NaHCO₃ ergogenic potential is associated with
318 its effects on dynamic buffering capacity³². High-intensity exercise produces an increase in the
319 rate of H⁺ accumulation, which can lead to intramuscular acidosis. Considering that acidosis has
320 been identified as a fatigue factor, intramuscular acidosis could interfere with several metabolic
321 and contractile processes, ultimately reducing force and power production associated with
322 fatigue during exercise¹. Thus, the main effect of NaHCO₃ supplementation seems to be related
323 to the increase of blood bicarbonate, which allows a greater efflux of H⁺ out of the active
324 muscle cells into the circulatory system. This idea aligns with previous studies where NaHCO₃
325 increases plasma lactate after repeated high-intensity efforts^{9,29}, and this study in which
326 NaHCO₃ was ingested before performing four Wingate tests interposed by 1.5 min of rest
327 among them.

328

329 The effect of caffeine on repeated sprint and power performance is mixed, potentially due
330 to some confounding variables such as inter-individual differences, caffeine adverse effects or
331 anxiety feelings among others². In an older study, Greer et al.³³ failed to report caffeine benefits
332 on power output during a 30-s high-intensity cycling bout using the Wingate test, similar to a
333 more recent study performed Duncan et al.³⁴ who examined the effects of acute caffeine
334 ingestion on upper- and lower-body Wingate test performance in twenty-two males, not
335 reporting significant findings when measuring lower-body W_{peak} and W_{mean} . However, Grgic³⁵
336 shows in his meta-analysis that caffeine ingestion stimulates an increase in W_{peak} and W_{mean}
337 during the Wingate test with a modest effect size of 0.27 (+4%) and 0.18 (+3%). Interestingly,
338 another study performed by Lee et al.³⁶ reported that caffeine ingestion enhanced sprint
339 performance involving a 90-s rest interval but did not benefit repeated sprints with a 20-s rest
340 interval, but that effect seems to be restricted to the first sprint since after repeated sprints total
341 work, best sprint or last sprint performance remained unchanged after acute caffeine
342 supplementation³⁷. This evidence seems to align with the result found in this study, where
343 caffeine intake stimulated an increase in time to reach W_{peak} in Wt1, but no other effect on
344 performance was detected.

345

346 The isolated and acute caffeine and NaHCO_3 intake stimulates ergogenic effects in high-
347 intensity and extenuating tasks. Both caffeine and NaHCO_3 have shown to delaying fatigue and
348 the muscular loss of ability to generate force and power^{3,4} through central-peripheral and
349 peripheral mechanism^{2,6-8}. Hence, it can be hypothesized that the co-ingestion of these
350 supplements may produce a synergic or additive effect since caffeine and NaHCO_3 may act
351 through similar (peripheral, Ca^{2+} bioavailability) or different mechanisms of action (central and
352 peripheral). However, in response to repeated sprints, the co-ingestion of NaHCO_3 and caffeine
353 did not produce a synergic or additive effect in peak or mean power production. Only it can be
354 argued that the decrease observed in time to reach W_{peak} in the first Wingate in the

355 NaHCO₃+CAF (-10.3%) could be a summative effect of the co-ingestion of both supplements
356 since isolated NaHCO₃ (-6.7%) and CAF (-8.5%) promote lower improvements in this
357 performance variable. The limited ergogenic effects produced by caffeine in this repeated sprint
358 protocol and a potential pharmacokinetic and pharmacodynamic interaction between
359 supplements may explain the absence of synergic effects when NaHCO₃ and caffeine are
360 acutely co-ingested. Thus, more studies are required to explore the potential synergic effects of
361 NaHCO₃ and caffeine in other types of exercise.

362 Previous studies reported side effects of NaHCO₃ and caffeine co-ingestion ⁵. In our
363 study, the side effects questionnaire revealed gastrointestinal discomfort after NaHCO₃
364 ingestion, alone or combined with caffeine, compared to placebo and caffeine despite the fact
365 that the bicarbonate intake was taken in two doses as recommended ⁹. Although most
366 participants did not report this effect, gastrointestinal discomfort could influence performance
367 in some participants limiting their repeated sprint performance.

368 No statistical differences were found in depression, anger, vigor, fatigue, confusion or
369 subjective vitality. Other studies found similar results after a 3-4 km cycling time trial in trained
370 cyclists ^{12,38}, four Wingate tests in the upper body ²⁵, or 2-3 bouts of a specific judo test ^{9,25}.
371 However, in tension, CAF increased compared to the rest of the supplementation protocols.
372 This finding is consistent with Jodra et al.²¹, who found a more pronounced effect of caffeine on
373 tension of elite athletes than recreational athletes after a single Wingate test. However, in our
374 study, no differences in vigor and subjective vitality were found, potentially due to the
375 strenuous exercise protocol followed in this study (1 vs 4 Wingates tests). Thus, although in
376 some cases caffeine, and potentially of sodium bicarbonate, effect or lack of effect may be
377 attributable to inter-individual differences caused by mood state or adverse effects ^{9,19}, this does
378 not seem to be the case in this study.

379 Finally, the major limitation of the present study was the impossibility of measuring
380 plasma levels of caffeine, bicarbonate and pH. This measurement would provide valuable

381 information regarding the absorption and effect of both supplements, as well as the potential
382 pharmacokinetic and pharmacodynamic interaction proposed.

383

384 **PRACTICAL APPLICATION**

385 This study shows that NaHCO₃ and caffeine co-ingestion did not cause a summative ergogenic
386 effect in repeated efforts. Therefore, based on the current results, we would recommend the
387 ingestion of NaHCO₃ and caffeine separately, rather than co-ingested, before any sport (e.g.,
388 cycling) that involves repeated sprints with a duration of ~30s each sprint. Thus, for athletes
389 seeking to improve performance in a single sprint would be useful the ingestion of 3 mg/kg of
390 caffeine, while for those athletes seeking to maintain their sprint performance after several
391 maximal efforts, the ingestion of 0.3 g/kg of NaHCO₃ would be desirable. Nonetheless, some
392 athletes may experience an increase in tension and muscular discomfort after caffeine intake or
393 gastrointestinal discomfort after NaHCO₃ intake. These potential health effects should be taken
394 into account in the development of individual nutritional supplement strategies.

395

396 **CONCLUSION**

397 The co-ingestion of NaHCO₃ and caffeine does not provide synergic effects after four Wingate
398 tests interposed by 1.5 min of rest, even though the isolated ingestion of caffeine increases the
399 first sprint performance and NaHCO₃ increases the third and fourth sprints performance.
400 Therefore, despite the potential existence of any pharmacokinetic and pharmacodynamic
401 interaction between supplements, acute co-ingestion of caffeine and NaHCO₃ does not produce
402 a synergic (central-peripheral or peripheral) effect on repeated sprint performance.

403

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409

410 **CONFLICT OF INTERESTS**

411 All the authors declare that they have no conflict of interest derived from the outcomes of this
412 study.

413

414 **AUTHOR CONTRIBUTIONS**

415 APL conceived the experiment. CF and APL designed the experiment. CF, PGE, DV and APL
416 collected the data. CF and APL analyzed and interpreted the data. CF, ALS and APL drafted the
417 manuscript. All authors read and approved the final version of the manuscript.

418

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542

543 **FIGURE LEGENDS**

544 **Figure 1.** Repeated sprint performance after the four supplementation protocols.

545 Peak power output (Fig. 1A), mean power output (Fig. 1B), time to reach W_{peak} (Fig. 1C),

546 Fatigue Index (Fig. 1D) and plasma lactate (Fig. 1E).

547 * P < 0.05 NaHCO₃ + CAF compared to PLA; # P < 0.05 NaHCO₃ compared PLA; \$, P < 0.05

548 CAF compared to PLA.

549 Abbreviations: W_{peak}, peak power output; NaHCO₃ + CAF, sodium bicarbonate plus caffeine;

550 NaHCO₃, sodium bicarbonate; CAF, caffeine; PLA, placebo.

551

552

553 **Supplementary figure 1.** Differences in repeated sprint performance after the four supplementation

554 protocols according to sex.

555 Peak power output in males and females (Suppl Fig. 1A and 1B), mean power output in males and

556 females (Suppl Fig. 1C and 1D), time to reach W_{peak} in males and females (Suppl Fig. 1E and 1F),

557 Fatigue Index in males and females (Suppl Fig. 1G and 1H) and plasma lactate in males and females

558 (Suppl Fig. 1I and 1J).

559 * P < 0.05 NaHCO₃ + CAF compared to PLA; # P < 0.05 NaHCO₃ compared PLA; \$, P < 0.05 CAF

560 compared to PLA.

561 Abbreviations: W_{peak}, peak power output; NaHCO₃ + CAF, sodium bicarbonate plus caffeine;

562 NaHCO₃, sodium bicarbonate; CAF, caffeine; PLA, placebo.

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