



## Article

# Isolated effects of caffeine and sodium bicarbonate ingestion on performance in the Yo-Yo test: A systematic review and meta-analysis

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## 1 **Abstract**

2 **OBJECTIVES:** To conduct a systematic review and a meta-analysis of studies exploring the  
3 effects of caffeine and/or sodium bicarbonate on performance in the Yo-Yo test.

4 **DESIGN:** Systematic review/meta-analysis.

5 **METHODS:** A total of six databases were searched, and random-effects meta-analyses were  
6 performed examining the isolated effects of caffeine and sodium bicarbonate on performance  
7 in the Yo-Yo test.

8 **RESULTS:** After reviewing 988 search records, 15 studies were included. For the effects of  
9 caffeine on performance in the Yo-Yo test, the meta-analysis indicated a significant favoring  
10 of caffeine as compared with the placebo conditions ( $p = 0.022$ ; standardized mean difference  
11 [SMD] = 0.17; 95% CI: 0.08, 0.32; +7.5%). Subgroup analyses indicated that the effects of  
12 caffeine were significant for the level 2 version of the Yo-Yo test, but not level 1. Four out of  
13 the five studies that explored the effects of sodium bicarbonate used the level 2 version of the  
14 Yo-Yo test. **The pooled SMD favored the sodium bicarbonate condition as compared with the  
15 placebo/control conditions ( $p = 0.007$ ; SMD: 0.36; 95% CI: 0.10, 0.63; +16.0%).**

16 **CONCLUSIONS:** This review demonstrates that isolated ingestion of caffeine and sodium  
17 bicarbonate enhances performance in the Yo-Yo test. Given these ergogenic effects, the  
18 intake of caffeine and sodium bicarbonate before the Yo-Yo test needs to be standardized  
19 **(i.e., either restricted or used in the same way before each testing session).** Furthermore, the  
20 results suggest that individuals competing in sports involving intermittent exercise may  
21 consider supplementing with caffeine or sodium bicarbonate for acute improvements in  
22 performance.

## 23 **Introduction**

24 The Yo-Yo intermittent recovery test was introduced in the 1990s and has gained substantial  
25 popularity as a method of estimating aerobic and anaerobic capacity in team-sports players.<sup>1</sup>  
26 In undertaking the Yo-Yo intermittent recovery test, participants are required to run distances  
27 of  $2 \times 20$  m at progressively increasing speeds. Each  $2 \times 20$  m work block is interspersed with  
28 a 10-s period of jogging around a marker placed 5 m behind the finish line. The test ends  
29 when the individual cannot complete the run within the prescribed time, on two consecutive  
30 occasions. The outcome of this test is the total covered distance. This test is comprised of two  
31 sub-levels, level 1 and level 2, with level 2 starting at a higher initial speed and necessitating a  
32 large contribution from the anaerobic energy system.<sup>1</sup> These tests are widely used in field  
33 settings as a practical method to: (a) determine current fitness status; and (b) prescribe  
34 training programs and explore their effectiveness.<sup>2</sup>

35  
36 Caffeine and sodium bicarbonate are two supplements that have been shown to acutely  
37 enhance exercise performance.<sup>3</sup> The effects of caffeine and sodium bicarbonate have also  
38 been explored in the context of Yo-Yo test performance, with equivocal findings.<sup>4-8</sup> Some  
39 reported an increase in performance following the ingestion of either caffeine or sodium  
40 bicarbonate, while others report that performance in this test is unaffected by the use of these  
41 supplements.<sup>4-8</sup> Differences between the studies such as the dose, ingestion timing, and  
42 training status may explain some of these discrepancies.<sup>9</sup> Perhaps even more importantly, the  
43 studies conducted on the effects of sports supplements tend to be performed with small  
44 sample sizes.<sup>3</sup> For example, one study that explored the effects of sodium bicarbonate on  
45 performance in the Yo-Yo test included only six participants.<sup>8</sup> Therefore, there remains a  
46 possibility that some studies were statistically underpowered to observe significant effects,  
47 which may have resulted in increased probabilities of type II errors. Meta-analysis is a

48 statistical method that can be used to overcome the limitation of underpowered studies  
49 because it allows combining of data from different cohorts to obtain a pooled estimated value.

50

51 To address the apparent discrepancies between individual studies two previous meta-analyses  
52 explored the effects of caffeine on performance in the Yo-Yo test. In the first analysis,  
53 Gonçalves Ribeiro et al.<sup>10</sup> reported no significant effects of caffeine on performance in this  
54 test. However, the analysis included only two studies with a combined number of participants  
55 amounting to 31. Such a small number of studies was included given that the authors limited  
56 their inclusion criteria to studies published between 2010 and 2015, even though no rationale  
57 was provided for this approach. Another limitation is that these authors included both cross-  
58 over and between-group study designs. This may be relevant given that the inter-individual  
59 variation in responses to caffeine ingestion is not as well controlled in between-group designs  
60 as in cross-over trials.<sup>11</sup>

61

62 In another review, Salinero et al.<sup>12</sup> reported that caffeine ingestion enhances performance in  
63 the Yo-Yo test by an effect size of 0.22 with a 95% confidence interval (CI) ranging from  
64 0.00 to 0.44. This meta-analysis included only four studies (pooled  $n = 57$ ) given that studies  
65 providing caffeine in doses lower than  $2 \text{ mg}\cdot\text{kg}^{-1}$  were not considered. Additionally, this  
66 review did not search through unpublished literature, which may have led to publication bias  
67 due to the “file drawer” syndrome. The “file drawer” syndrome suggests that studies with  
68 larger effect sizes are more likely to be published than those with small or non-significant  
69 effect sizes; therefore, including only published studies may present a source of bias in a  
70 given meta-analysis.<sup>13, 14</sup> Additionally, two studies<sup>4, 15</sup> were not included in the review by  
71 Salinero et al.<sup>12</sup> and new studies on this topic have been published since this review.<sup>7, 16</sup> All  
72 these studies generated data for over 100 additional participants. Therefore, an updated meta-

73 analysis that would include these studies could contain almost a threefold increase in the  
74 pooled sample size from the most recent meta-analysis<sup>12</sup> thus increasing the veracity of the  
75 findings.

76

77 While the effects of caffeine have been explored using a meta-analytic approach, no previous  
78 meta-analyses explored the effects of sodium bicarbonate on performance in the Yo-Yo test.  
79 A meta-analysis exploring the effects of sodium bicarbonate on performance in this test is  
80 needed because: (a) the equivocal evidence on this topic presented in the literature;<sup>5,6,8</sup> and,  
81 (b) the common anecdotal use of sodium bicarbonate by athletes.<sup>17</sup> Therefore, the aim of this  
82 review was to explore the effects of caffeine and sodium bicarbonate on performance in the  
83 Yo-Yo testing using a meta-analysis and highlight the practical application of these findings.

84

## 85 **Methods**

86 For this review, peer-reviewed literature in the form of journal articles and unpublished  
87 literature in the form of a thesis, dissertations, or conference abstracts that investigated the  
88 effects of caffeine and/or sodium bicarbonate ingestion on performance in the Yo-Yo test was  
89 examined. The search for studies was conducted on April 14th, 2019. The following databases  
90 were searched: ERIC, PubMed/MEDLINE, SPORTDiscus, Open Access Thesis and  
91 Dissertations, Web of Science, and Scopus. Additionally, searches were performed within  
92 ResearchGate. In all of these databases, the following search syntax was used: ("caffeine" OR  
93 "supplement" OR "coffee" OR "ergogenic" OR "NaHCO<sub>3</sub>" OR "sodium bicarbonate") AND  
94 ("yo-yo" OR "yoyo" OR "yo yo"). Secondary searches were performed by screening the  
95 reference lists of the included studies, and by exploring the papers that cited the included  
96 studies. The search was performed independently by two authors (JG and AG).

97

98 To be included in the review, studies were required to satisfy the following criteria: (1)  
99 published in English; (2) included apparently healthy humans as participants; and (3)  
100 employed a cross-over study design and explored the acute effects of caffeine and/or sodium  
101 bicarbonate on performance in any variant of the Yo-Yo test.

102

103 Two authors of the review (JG and AG) extracted the following data from the included  
104 studies: (1) author names and publication year; (2) sample size and the sample characteristics;  
105 (3) the variant of the Yo-Yo test used for the testing; and (4) main findings regarding the  
106 effects of caffeine and/or sodium bicarbonate on performance. In cases where the required  
107 data was presented in figures, the Web Plot Digitizer software was used for the extraction of  
108 raw values.

109

110 The PEDro checklist was used for assessing the quality of the included studies.<sup>18</sup> This  
111 checklist has 11 items; these items refer to eligibility criteria, randomization and blinding of  
112 participants, blinding of assessors, the number of participants completing all testing sessions,  
113 and reporting of data for the key outcomes. While this checklist has 11 points, the first item is  
114 not included in the summary score, and therefore, the maximum number of points is 10.

115 Based on the summary scores, the studies were classified as being of excellent quality (9-10  
116 points), good quality (6-8 points), fair quality (4-5 points) and poor methodological quality  
117 (<3 points).<sup>19-21</sup> The quality assessment was conducted independently by two authors of the  
118 review (JG and AG).

119

120 The mean  $\pm$  standard deviation performance data extracted from the included studies were  
121 converted to standardized mean differences (SMDs) and their respective 95% CIs. The  
122 following data are required to calculate SMDs: (1) Yo-Yo test performance mean  $\pm$  standard

123 deviation of the caffeine/sodium bicarbonate and placebo/control trials, (2) total sample size,  
124 and (3) inter-trial correlation. Inter-trial correlation was not presented in any of the included  
125 studies. As suggested in the Cochrane Handbook the following formula was used to estimate  
126 the correlation:

127

$$128 \quad r = \frac{S_{\text{placebo/control}}^2 + S_{\text{treatment}}^2 - S_D^2}{2 \cdot S_{\text{placebo/control}} \cdot S_{\text{treatment}}}$$

129

130

131  $S$  represents the standard deviation while  $S_D$  is the standard deviation of the difference score,  
132 which was calculated as:

133

$$134 \quad S_D = \left( \frac{S_{\text{placebo/control}}^2}{n} + \frac{S_{\text{treatment}}^2}{n} \right)^{1/2}$$

135

136 One of the studies that explored the effects of caffeine used multiple caffeine doses; for this  
137 study, SMDs and variances were calculated for each dose separately and the average values  
138 were used for the analysis. Two meta-analyses were performed: (1) for the effects of caffeine  
139 on performance in the Yo-Yo test; and (2) for the effects of sodium bicarbonate on  
140 performance in the Yo-Yo test. In the meta-analysis that focused on the effects of caffeine, a  
141 sensitivity analysis was performed by excluding one study in which caffeine was not ingested  
142 as it was provided in a mouth rinsing form.<sup>22</sup> A subgroup analysis was performed for studies  
143 exploring the effects of caffeine on Yo-Yo intermittent recovery level 1 and for those  
144 exploring the level 2 variant of the test. In the meta-analysis for the effects of sodium  
145 bicarbonate, a sensitivity analysis was performed by excluding the only study that used the

146 level 1 version of the test. SMD values of <0.20, 0.20-0.39, 0.40-0.59, 0.60-0.80, and >0.80  
147 were considered to represent trivial, small, medium, large, and very large effects, respectively.  
148 In each analysis, the  $I^2$  statistic was used to explore heterogeneity with  $I^2$  values of <50%, 50  
149 to 75%, and >75% considered as low levels, moderate levels, and high levels of  
150 heterogeneity.

151

152 In addition to 95% CIs, 95% prediction intervals (95% PI) were calculated for both analyses  
153 by using the number of included studies in the meta-analysis, the pooled SMD, the upper limit  
154 of the 95% CI and the tau-squared values. The 95% PI denotes the range in which the effect  
155 size of a future study conducted on the topic will most likely be. Funnel plots asymmetry was  
156 explored only for the effects of caffeine given that there were less than 10 studies included in  
157 the analyses for sodium bicarbonate. Percent differences between supplement ingestion  
158 conditions and the placebo/control conditions were also calculated. The random-effects model  
159 was used for both analyses. The statistical significance threshold was set at  $p < 0.05$ . All  
160 analyses were performed using the Comprehensive Meta-analysis software, version 2 (Biostat  
161 Inc., Englewood, NJ, USA).

162

## 163 **Results**

164 The total number of search results across all databases was 164. Of this number of search  
165 results, 21 full-text papers were read and 12 studies were included. The remaining documents  
166 were excluded based on the title or abstract. The secondary searches resulted in another 824  
167 search results and in the inclusion of three additional studies. Therefore, in total, 15 studies  
168 were included; 13 studies were published as full-text manuscripts in peer-reviewed journals,  
169 one study was published as a conference abstract, and one study is a part of a master's thesis.<sup>4-</sup>  
170 <sup>8,13,15,16,22-28</sup> Eleven studies explored the effects of caffeine, whereas five studies examined the



171 effects of sodium bicarbonate on Yo-Yo test performance (Table 1). One study included  
172 examined the isolated effects of both caffeine and sodium bicarbonate.

173

174 For studies that explored the effects of caffeine, the pooled number of participants across all  
175 studies was 156 (21 females). The average sample size per study was 14 participants. In all of  
176 the included studies, the sample was comprised of athletes. The doses of caffeine in the  
177 studies ranged from 1 mg·kg<sup>-1</sup> to 6 mg·kg<sup>-1</sup>. Five studies provided absolute doses of caffeine  
178 ranging from 200 to 500 mg. One study used a caffeine mouth rinsing form where 1.2% of the  
179 25 ml solution was caffeine. Timing of caffeine administration before exercise ranged from  
180 immediately before (one study), 5 minutes (two studies), 45 minutes (two studies), 50 minutes  
181 (one study), 60 minutes (four studies) and 70 minutes (one study) before exercise. Five  
182 studies used the intermittent Yo-Yo recovery test level 1 and six used the level 2 version.

183

184 The pooled number of participants for the studies that explored the effects of sodium  
185 bicarbonate was 46 (all males). The average sample size per study was 9 participants. Three  
186 studies included athletes as their study participants while two included recreationally active  
187 individuals. The doses of sodium bicarbonate in the studies ranged from 0.2 g·kg<sup>-1</sup> to 0.4 g·kg<sup>-1</sup>.  
188 <sup>1</sup>. Sodium bicarbonate was ingested 40 minutes pre-exercise (one study), 60 minutes (one  
189 study), and 90 minutes pre-exercise (two studies). One study used a protocol that included  
190 splitting up a dose of 0.4 g·kg<sup>-1</sup> into five smaller doses taken at 90, 80, 70, 60, and 50 minutes  
191 pre-exercise. Four studies used the intermitted Yo-Yo recovery test level 2, with only one  
192 using the level 1 version.

193

194 Out of the eleven studies that explored the effects of caffeine on Yo-Yo test performance,  
195 eight were classified as being of excellent quality with two studies being classified as good

196 methodological quality and one as fair methodological quality. Out of the five studies that  
197 explored the effects of sodium bicarbonate on Yo-Yo test performance, three were classified  
198 as being of excellent quality, one as good methodological quality and one study as being of  
199 fair methodological quality. No studies were classified as being poor methodological quality.  
200 Individual scores for the quality assessment can be found in Supplementary file 1.

201

202 The meta-analysis for the effects of caffeine included a total of ten studies given that in one  
203 study, the data required for the analysis was not presented and the authors did not provide the  
204 data upon written request.<sup>22</sup> The main meta-analysis indicated a significant difference ( $p =$   
205  $0.022$ ) between the caffeine and placebo conditions with the SMD favoring the caffeine  
206 condition (SMD: 0.17; 95% CI: 0.08, 0.32; percent change: +7.5%;  $I^2$ : 28%; 95% PI: -0.32,  
207 0.66; Figure 1). No funnel plot asymmetry was observed. In the sensitivity analysis in which  
208 the study that provided caffeine in a mouth rinsing form was excluded, the SMD values  
209 increased to 0.20 (95% CI: 0.05, 0.36;  $p = 0.009$ ; percent change: +8.5%;  $I^2$ : 26%). In the  
210 subgroup analysis for the level 1 Yo-Yo test the SMD was 0.02 (95% CI: -0.21, 0.25;  $p =$   
211  $0.880$ ; percent change: +2.5%;  $I^2$ : 0%). In the subgroup analysis for the level 2 Yo-Yo test the  
212 SMD was 0.31 (95% CI: 0.12, 0.51;  $p = 0.002$ ; percent change: +14.4%;  $I^2$ : 11%).

213

214 The meta-analysis for the effects of sodium bicarbonate indicated a significant difference ( $p =$   
215  $0.007$ ) between the sodium bicarbonate and placebo/control conditions. The pooled SMD  
216 favored the sodium bicarbonate condition (SMD: 0.36; 95% CI: 0.10, 0.63; percent change:  
217 +16.0%;  $I^2$ : 14%; 95% PI: -0.61, 1.33; Figure 2). In the sensitivity analysis in which the study  
218 that used the level 1 Yo-Yo test version was excluded, the SMD values increased to 0.39  
219 (95% CI: 0.08, 0.70;  $p = 0.013$ ; percent change: +17.5%;  $I^2$ : 25%).

220

## 221 **Discussion**

222 This review reports that isolated ingestion of caffeine and sodium bicarbonate enhances  
223 performance in the Yo-Yo test. Both supplements seem to produce moderate performance-  
224 enhancing effects. Due to these acute ergogenic effects, the intake of caffeine and sodium  
225 bicarbonate before the Yo-Yo test needs to be standardized. The results also suggest that  
226 individuals competing in sports involving intermittent exercise may consider supplementing  
227 with caffeine or sodium bicarbonate for acute improvements in performance.

228

229 This meta-analysis adds further evidence that caffeine ingestion enhances performance in  
230 field-based tests of fitness. These results are in line with prior work in the area, most of which  
231 is based on tests performed in the laboratory.<sup>14</sup> Caffeine's ergogenic effect is likely related to  
232 its binding to adenosine receptors.<sup>29</sup> Caffeine has a similar structure to adenosine and  
233 therefore, when ingested, caffeine binds to A<sub>1</sub> and A<sub>2A</sub> receptors, ultimately blunting the  
234 fatiguing effects of adenosine. As a result, acute caffeine ingestion may reduce perceived  
235 effort and increase physical performance. Caffeine ingestion may also enhance motor unit  
236 recruitment, thus leading to more forceful muscle contractions.<sup>30,31</sup> These mechanisms might  
237 explain why the pooled SMD increased when the study that provided caffeine in the mouth  
238 rinsing form was excluded.<sup>22</sup> After excluding the study that utilized caffeine mouth rinsing,  
239 the SMD increased from 0.17 to 0.20. Specifically, caffeine provided in this form does not  
240 increase plasma caffeine concentration<sup>32</sup> which is a likely prerequisite for an ergogenic effect  
241 of caffeine.

242

243 Due to a small number of included studies, previous meta-analyses did not examine if the  
244 effects of caffeine differ between level 1 and level 2 versions of the Yo-Yo test. These

245 subgroup analyses were performed in the present review and indicated that the effects of  
246 caffeine were significant only for the level 2 version. Such findings may suggest that caffeine  
247 is more effective in tests with a greater contribution from the anaerobic energy system.<sup>1</sup> These  
248 results were obtained from studies that examined the effects of caffeine on either level 1 or  
249 level 2 versions of the test. However, given the inter-individual variation in responses to  
250 caffeine ingestion,<sup>11,33,34</sup> future studies may consider comparing the effects of caffeine on  
251 performance in the level 1 and level 2 version of the test in the same group of participants.

252

253 Based on the results of this review, sodium bicarbonate is effective for acute increases in Yo-  
254 Yo test performance. Sodium bicarbonate may elicit its ergogenic effects by: (a) acutely  
255 increasing blood bicarbonate and leading to a greater efflux of hydrogen ions ( $H^+$ ) and lactate  
256 out of the active muscles and into the circulation; and subsequently, (b) maintenance of  
257 intramuscular pH.<sup>35,36</sup> Following acute sodium bicarbonate ingestion and subsequent increase  
258 in blood bicarbonate concentration, the resultant pH gradient between the intracellular and  
259 extracellular environments favors the efflux of  $H^+$  from the exercising muscle to blood, aiding  
260 intracellular pH regulation and reducing fatigue.<sup>35,36</sup>

261

262 While no previous meta-analysis examined the effects of sodium bicarbonate on Yo-Yo test  
263 performance, other meta-analyses have focused on different aspects of exercise performance.  
264 For example, Matson and Tran<sup>37</sup> combined studies that explored the effects of sodium  
265 bicarbonate on various exercise tests, with some lasting over 30 minutes and others being of  
266 very short duration, and maximal intensity (e.g., 10-second “all-out” sprints). The pooled  
267 ergogenic effect size of sodium bicarbonate was 0.44. The magnitude of the effect is  
268 comparable to the effect size observed herein, even though an argument can be made that the

269 pooling of vastly different exercise tests (with different physiological demands) in the  
270 analysis by Matson and Tran<sup>37</sup> might have been a methodological limitation. A meta-analysis  
271 by Carr et al.<sup>38</sup> reported an ergogenic effect of acute sodium bicarbonate ingestion that  
272 amounted to 1.7%. The performance-enhancing effect in the analysis by Carr et al.<sup>38</sup> was  
273 small likely because it included many of the older studies that used protocols in which all  
274 participants ingest sodium bicarbonate around 60 to 90 minutes before exercise.<sup>9,39,40</sup>  
275 However, in recent years, studies have started using individualized time to peak blood  
276 bicarbonate protocols, and they generally report greater effect sizes.<sup>9,39,40</sup> The majority of  
277 studies included in the meta-analysis utilized performance tests lasting up to 30 seconds (i.e.,  
278 much shorter than the average duration of the Yo-Yo test), which limits further comparison of  
279 the results. Nonetheless, this review reinforces the suggestion of the International Olympic  
280 Committee that acute sodium bicarbonate ingestion enhances short-term high-intensity  
281 exercise performance.<sup>3</sup>

282

283 The results presented in this review highlight the need for standardizing caffeine and sodium  
284 bicarbonate intake before the Yo-Yo test. **In other words, supplementation with caffeine or  
285 sodium bicarbonate should be either restricted or used in the same way before each testing  
286 session. If their use is not standardized, some individuals may ingest caffeine or sodium  
287 bicarbonate before the testing and possibly experience a supplement-induced improvement  
288 performance in the Yo-Yo test.** This may be especially important to control when using the  
289 Yo-Yo test for the evaluation of the effectiveness of a given training program. Additionally,  
290 standardization of caffeine and sodium bicarbonate intake may be important for studies that  
291 focus on the reliability of the Yo-Yo test.<sup>41</sup> If not standardized, ingestion of these supplements  
292 may lead to improvements in performance by small to moderate effects (i.e., pooled SMDs in

293 this review ranged from 0.17 to 0.39); if not standardized, this improvement may affect the  
294 validity of the data and lead to incorrect interpretation of the results.

295

296 One additional value of these substances is that their ingestion may improve performance in  
297 training sessions or competitions with similar energy demands as the Yo-Yo test. This may be  
298 especially relevant in soccer given that Krustup et al.<sup>42</sup> observed that the performance in the  
299 Yo-Yo test is significantly correlated ( $r=0.81$ ) with the number of high-intensity running  
300 actions performed at the end of each half of a game. These results suggest that caffeine or  
301 sodium bicarbonate ingestion may even improve performance directly during sports  
302 competitions. These supplements may also have the potential to enhance training responses  
303 and adaptations; albeit, future long-term studies are needed to establish such effects. Still,  
304 while the Yo-Yo test is valid for determining an individual's capacity to perform repeated  
305 exercise,<sup>1</sup> future studies may consider exploring the effects of caffeine and sodium  
306 bicarbonate using more specific team sport stimulations. **For example, one study explored the  
307 effects of caffeine while using a performance test simulating physical and skill demands of a  
308 rugby union game which included seven circuits in each of two 40-min halves with a 10-min  
309 half-time rest.**<sup>43</sup>

310

311 Using the PEDro checklist, the included studies are generally sound from a methodological  
312 quality perspective. Nonetheless, several included studies used a single-blind protocol which  
313 offers evidence of lower that the use of the "gold standard" double-blind study design. Of the  
314 studies that blinded the participants, only two<sup>16,28</sup> explored the effectiveness of this blinding  
315 by asking the participants to indicate which condition was the placebo and which the  
316 caffeine/sodium bicarbonate one. This limitation needs to be addressed in future studies given

317 that correct supplement identification may impact exercise outcomes and therefore lead to  
318 bias in the results.<sup>44</sup>

319

320 The main limitation is that some of the studies included in the meta-analysis on the effects of  
321 sodium bicarbonate used a placebo condition as the comparison and other employed a non-  
322 supplement, control trial. This methodological aspect may have affected the results given that  
323 the act of ingesting a capsule (even though it does not contain an ergogenic compound) can  
324 produce improvements in performance due to the placebo effect. However, the largest  
325 improvement in performance following sodium bicarbonate ingestion was in a study<sup>26</sup> that  
326 used a placebo vs. sodium bicarbonate comparison (SMD: 0.93; +30%) which may suggest  
327 that this limitation may not be particularly impactful in this specific context (even though it  
328 needs to be stated). Additionally, it needs to be mentioned that only one<sup>25</sup> of the five studies  
329 that explored the effects of sodium bicarbonate used a sodium-matched placebo comparison.  
330 Given that there are cases in which sodium can also be ergogenic,<sup>45</sup> future studies on this  
331 topic should consider adding a condition with an equimolar amount of salt to the sodium  
332 bicarbonate condition to isolate the effects of bicarbonate.<sup>46</sup>

333

334 In the included studies, caffeine and sodium bicarbonate were provided in isolation. Given  
335 that both supplements appear to be ergogenic, future work is needed to explore whether their  
336 combined ingestion provides any additive benefits. It is possible that the combination of these  
337 supplements would provide even greater effects because acute ingestion of caffeine and  
338 sodium bicarbonate enhances performance through different physiological mechanisms. As  
339 stated previously, caffeine's ergogenic effects are explained by its tendency to bind to  
340 adenosine receptors while sodium bicarbonate elicits its ergogenic effect through pH

341 regulation. While there are several studies conducted on this topic, future work is needed to  
342 provide greater clarity on the issue of supplement interactions.<sup>35</sup> This area may be particularly  
343 relevant for athletes given that athletes often ingest more than one supplement,<sup>47</sup> and that  
344 caffeine is often ingested as part of a multi-ingredient pre-workout supplement, or energy  
345 drink. Future studies are needed to explore optimal doses of caffeine and sodium bicarbonate  
346 that have the largest effect on performance while producing the least side-effects.<sup>48</sup> In  
347 addition to doses, future work is need on the optimal timing of ingestion. Timing of ingestion  
348 may be especially important to investigate when it comes to sodium bicarbonate given that  
349 there is very large inter-individual variability in responses to sodium bicarbonate  
350 ingestion,<sup>9,39,40</sup> and the timing of ingestion is also affected by the mode of delivery.<sup>49</sup>

351

## 352 **Conclusion**

353 The results of the present review indicate that isolated caffeine and sodium bicarbonate  
354 ingestion enhances performance in the Yo-Yo test. Given these acute performance-enhancing  
355 effects, the intake of caffeine and sodium bicarbonate before the Yo-Yo test needs to be  
356 standardized (*i.e., either restricted or used in the same way before each testing session*). Also,  
357 the results suggest that these substances may improve performance during exercise tasks with  
358 similar energy demands to the Yo-Yo test and athletes competing in intermittent sports may  
359 consider supplementing with caffeine or sodium bicarbonate for acute increases in  
360 performance.



361 **Conflicts of interest**

362 None.

363 **References**

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