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**Title:** Sodium bicarbonate improves 4 km time trial cycling performance when individualised to time to peak blood bicarbonate in trained male cyclists.

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#### Abstract

The aim of this study was to investigate the effects of sodium bicarbonate (NaHCO<sub>3</sub>) on 4 km cycling time trial (TT) performance when individualised to a predetermined time to peak blood bicarbonate (HCO<sub>3</sub><sup>-</sup>). Eleven male trained cyclists volunteered for this study (height 1.82  $\pm$  0.80 m, body mass (BM) 86.4  $\pm$  12.9 kg, age 32  $\pm$  9 years, peak power output (PPO) 382  $\pm$  22 W). Two trials were initially conducted to identify time to peak HCO<sub>3</sub><sup>-</sup> following both 0.2 g kg<sup>-1</sup> BM (SBC2) and 0.3 g kg<sup>-1</sup> BM (SBC3) NaHCO<sub>3</sub>. Thereafter, on three separate occasions using a randomized, double-blind, crossover design, participants completed a 4 km TT following ingestion of either SBC2, SBC3, or a taste-matched placebo (PLA) containing 0.07 g kg<sup>-1</sup> BM sodium chloride (NaCl) at the predetermined individual time to peak HCO<sub>3</sub><sup>-</sup>. Both SBC2 (-8.3  $\pm$  3.5 s; p <0.001, *d* =0.64) and SBC3 (-8.6  $\pm$  5.4 s; p =0.003, *d* =0.66) reduced the time to complete the 4 km TT, with no difference between SBC conditions (mean difference = 0.2  $\pm$  0.2 s; p =0.87, *d* =0.02). These findings suggest trained cyclists may benefit from individualising NaHCO<sub>3</sub> ingestion to time to peak HCO<sub>3</sub><sup>-</sup> to enhance 4 km TT performance. **Key words:** buffering, metabolic alkalosis, dosage, individual pursuit

# 1 Introduction

2 Competitive cycling is reflective of high-intensity exercise, particularly in events such as the 3 individual and team pursuit, which entails completion of a 4 km time trial (TT). The typical 4 duration of this event ranges between 4 (world record times) and 7 min (recreational riders), 5 and because of this, a large energy supply is provided by anaerobic glycolysis (Gastin, 2001). 6 With such a demand an exponential accumulation of metabolites including inorganic 7 phosphate, hydrogen ions (H<sup>+</sup>), and lactate occurs (Westerblad et al., 2002; Allen et al., 2008). 8 Due to the inverse relationship between H<sup>+</sup> and pH, this process causes metabolic acidosis and 9 results in a decrease in blood and muscle pH (Allen et al., 2008). Whilst there is no singular 10 mechanism of peripheral fatigue, perturbations to acid base balance have been implicated to 11 inhibit enzyme activity (e.g. glycogen phosphorylase) and calcium ion (Ca<sup>2+</sup>) cross-bridge 12 binding (Fitts, 2008, 2016). Preventative strategies such as the ingestion of nutritional 13 ergogenic aids may therefore be beneficial to mitigate such local acid-base disturbances in 14 active musculature (Christensen, Shirai, Ritz, & Nordsborg, 2017; Matson & Tran, 1993).

15

16 Ingestion of sodium bicarbonate (NaHCO<sub>3</sub>), a known buffering agent, can reinforce acid base 17 balance by producing a state of metabolic alkalosis (increased pH and HCO<sub>3</sub><sup>-</sup>) (McNamara & 18 Worthley, 2001). Increases in pH typically result in a greater efflux of H<sup>+</sup> and lactate from 19 active musculature into extracellular compartments, due to a greater intra-extracellular 20 gradient, whilst elevated HCO<sub>3</sub><sup>-</sup> can be utilised to buffer against H<sup>+</sup> within extracellular 21 compartments (Bishop, Edge, Davis and Goodman, 2004). The resulting effect is more work 22 completed during exercise of high intensities, which in turn, will improve exercise capacity or 23 performance (Bishop et al., 2004; Marx et al., 2002). It is therefore important to heighten the 24 level of blood alkalosis via changes in pH and HCO<sub>3</sub><sup>-</sup> prior to exercise (Gough, Deb, Sparks & 25 McNaughton, 2017a; Jones et al., 2016). Common practice is to prescribe NaHCO<sub>3</sub> between a 26 set time of between 60 and 90 mins for all participants (Carr, Hopkins and Gore, 2011; Price 27 and Singh, 2008; Siegler et al., 2009). In a recent study, however, it was reported time to peak HCO<sub>3</sub><sup>-</sup> occurred between 40 and 125 min (Gough et al., 2017a), with a similar variation 28 29 observed in other dose-response studies (Jones et al., 2016; Miller et al., 2016). Many 30 participants may not therefore achieve peak alkalosis at the start of exercise, which might 31 explain, in part, the lack of an ergogenic effect of NaHCO<sub>3</sub> supplemented at 100 min (Correia-32 Oliveira et al., 2017) and 150 min (Callahan, Parr, Hawley & Burke, 2017) in other 4 km 33 cycling TT studies.

34

35 In response to such variation in time to peak alkalosis it is recommended that either time to 36 peak pH or HCO<sub>3</sub><sup>-</sup> is predetermined prior to use for an exercise bout, as this accounts for the 37 inter-individual variation commonly observed (McNaughton et al., 2016; Miller et al., 2016; 38 Jones et al., 2016; Gough et al., 2017c). Indeed, preliminary studies to date have displayed 39 ergogenic benefits of NaHCO<sub>3</sub> individualised to a predetermined peak pH in cycling 40 performance (Miller et al., 2016; Deb et al., 2017). Gough et al. (2017a) however, recently 41 demonstrated greater reliability of time to peak HCO3<sup>-</sup> compared to time to peak pH with 42 Intraclass Correlation Coefficient (ICC) analysis (r = 0.94 vs. 0.71). It may therefore be more 43 appropriate to determine the effects of NaHCO<sub>3</sub> on HCO<sub>3</sub><sup>-</sup> responses, particularly if the athlete 44 wishes to achieve peak alkalosis consistently. Nonetheless, no study to date has investigated 45 the potential ergogenic effects of NaHCO<sub>3</sub> supplementation determined by a predetermined 46 individual time to peak  $HCO_3^-$  on an exercise protocol reflective of competitive cycling such 47 as a 4 km TT.

48

Investigations into the ergogenic effects of individualising NaHCO<sub>3</sub> to a predetermined time
to peak pH have prescribed an amount of 0.3 g·kg<sup>-1</sup> BM (Miller et al., 2016; Deb et al., 2017).

51 This is likely due to early research by McNaughton (1992) reporting a dose-dependent effect 52 on performance, with 0.3 gkg<sup>-1</sup> BM NaHCO<sub>3</sub> improving total work done (TWD) to a greater magnitude than 0.2 gkg<sup>-1</sup> during 60 s of maximal cycling; whilst meta-analyses have also 53 shown a meaningful effect on exercise performance following 0.3 gkg<sup>-1</sup> BM NaHCO<sub>3</sub> (Peart 54 55 et al., 2012; Carr et al., 2011). Despite this, there is a paucity of literature investigating the 56 dose-dependent ergogenic effects from smaller doses of NaHCO<sub>3</sub> on exercise performance. The greater magnitude of effect between 0.3 gkg<sup>-1</sup> and 0.2 gkg<sup>-1</sup> BM NaHCO<sub>3</sub> reported by 57 McNaughton (1992) for instance, was non-significant and only considered one exercise 58 59 duration/intensity and participant cohort (recreationally active). Furthermore, McKenzie, 60 Coutts, Stirling, Hoeben and Kuzara (1986) reported a negligible 0.3% difference between 0.15 61 g·kg<sup>-1</sup> BM and 0.3 g·kg<sup>-1</sup> BM NaHCO<sub>3</sub> in a cycling time to volitional exhaustion test at 125% VO<sub>2max</sub>. Based on such limited evidence, further research is warranted exploring the dose-62 63 dependent effects of NaHCO<sub>3</sub>.

64

A further concern of a 0.3 g kg<sup>-1</sup> BM NaHCO<sub>3</sub> ingestion strategy is the commonly reported 65 66 gastrointestinal (GI) discomfort symptoms such as stomach cramp, diarrhoea, and in extreme 67 cases, vomiting, which can have major negative implications for exercise performance 68 (Saunders et al., 2014; Gough et al., 2017a, 2017b). It is therefore important to maximise the 69 potential ergogenic effect through attaining peak buffering capacity, whilst also managing the severity of (GI) discomfort. Given that smaller amounts of NaHCO<sub>3</sub> (i.e. 0.2 gkg<sup>-1</sup> BM) are 70 71 associated with lower instances and severity of GI discomfort (Gough et al., 2017a, 2017c), it 72 may be prudent to suggest this amount is a better option practically to the athlete aiming to 73 enhance their performance, as long as ergogenic benefits are still evident.

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75 To heighten the likeliness of an ergogenic benefit and mitigate the severity of GI discomfort, 76 0.2 g kg<sup>-1</sup> BM NaHCO<sub>3</sub> individualised to a predetermined time to peak HCO<sub>3</sub><sup>-</sup> may be suitable. Gough et al. (2017a) reported a  $5.7 \pm 0.9$  mmol<sup>-1</sup> increase of HCO<sub>3</sub><sup>-</sup> following 0.2 gkg<sup>-1</sup> BM 77 78 NaHCO<sub>3</sub> using a time to peak HCO<sub>3</sub><sup>-</sup> strategy, which is superior to the  $3.9 \pm 0.9$  mmol·l<sup>-1</sup> mean 79 change reported in a meta-analysis following a standardised 0.3 g kg<sup>-1</sup> BM NaHCO<sub>3</sub> dose (Carr et al., 2011). These changes in acid base balance following 0.2 gkg<sup>-1</sup> BM NaHCO<sub>3</sub> are also 80 81 close to the 6 mmol<sup>-1</sup> increase purported to lead to an ergogenic effect on performance (Matson & Tran, 1993; Jones et al., 2016). These data combined, suggest 0.2 gkg<sup>-1</sup> BM 82 83 NaHCO<sub>3</sub> individualised to a pre-determined time to peak HCO<sub>3</sub><sup>-</sup> achieves the required acid 84 base balance changes that may improve performance, whilst also reducing the symptoms of GI 85 discomfort. Despite this, no literature to date has investigated the dose-dependent effects (i.e. 0.2 gkg<sup>-1</sup> vs. 0.3 gkg<sup>-1</sup> BM NaHCO<sub>3</sub>) on exercise performance when individualised to a 86 87 predetermined time to peak  $HCO_3^{-}$ . The purpose of this study, therefore, was to investigate the effects of both 0.2 gkg<sup>-1</sup> BM (SBC2) and 0.3 gkg<sup>-1</sup> BM (SBC3) NaHCO<sub>3</sub> individualised to a 88 89 predetermined time to peak HCO<sub>3</sub><sup>-</sup> on 4 km TT performance. We hypothesised that both SBC2 90 and SBC3 would reduce the time required to complete the 4 km TT.

91

#### 92 Materials and Methods

#### 93 Participants

A priori power calculation conducted using SPSS Sample Power 3 (IBM, Chicago, IL, USA) displayed a sample size of 11 would allow detection of a 3 s change with high statistical power  $(\beta = 0.80; 0.05 = \alpha$  level). This set criterion was used to detect a difference between NaHCO<sub>3</sub> treatments (i.e. SBC2 vs. SBC3) and between SBC treatments and the placebo, as this is the typical difference required to determine medal positions for the men's individual pursuit and similar events at Olympic Games (Christensen et al., 2017). Eleven male trained cyclists 100 therefore volunteered for this study (height  $1.82 \pm 0.8$  m, body mass  $86.4 \pm 12.9$  kg, age  $32 \pm$ 101 9 years, peak power output (PPO)  $382 \pm 22$  W) with a weekly training frequency of  $\geq 3$  times, 102 for a total of  $\geq$ 5 hours per week, and for a minimum of 2 years training experience, which was 103 specifically in cycling. Based on these descriptors, participants met the criteria of 'trained 104 cyclist' as described by De Pauw et al. (2013). Participants were also excluded if they had 105 ingested any nutritional buffers (such as beta alanine) in the prior 6 months of the study. Ethical 106 approval was obtained from the Departmental Research Ethics Committee and each participant 107 provided written informed consent prior to experimental testing.

108

### 109 Experimental overview

110 Participants visited the laboratory on six occasions in a randomised, crossover and double blind 111 designed study (2 x identification of peak blood HCO<sub>3</sub>-, 3 x cycling TT's). Constraints on 112 ingestion of alcohol and participation in any strenuous/unaccustomed exercise were in place 113 24 hours prior to each trial. Caffeine was also prohibited 12 hours prior to any trial. Written 114 logs of nutritional intake were taken, with intake from the first trial replicated for subsequent 115 trials. Participants visited the laboratory in a four-hour postprandial state and trials were 116 conducted at the same time of day to account for circadian rhythms (Reilly, 1990). 117 Experimental trials were separated by at least three days to allow acid base balance variables 118 to return to normal resting concentrations (Siegler et al., 2009).

119

# 120 Identification of time to peak blood bicarbonate

121 On two separate occasions participants ingested either 0.2 g·kg<sup>-1</sup> BM NaHCO<sub>3</sub> (SBC2) or 0.3 122 g·kg<sup>-1</sup> BM NaHCO<sub>3</sub> (SBC3) mixed with 400 ml of water and 50 ml double strength and sugar-123 free blackcurrant cordial to identify time to peak blood HCO<sub>3</sub><sup>-</sup> and pH. Whilst quietly resting 124 and seated, finger prick capillary blood samples were collected in a 100µl sodium heparin125 coated glass clinitube every 10 min for analysis of blood HCO<sub>3</sub><sup>-</sup> and pH over a 120 min period 126 using a blood gas analyser (ABL800 BASIC, Radiometer Medical Ltd. Denmark). The highest 127 HCO<sub>3</sub><sup>-</sup> value was used as a determination of time to peak HCO<sub>3</sub><sup>-</sup> and this determined the timing 128 of ingestion for experimental trials. Supplementation of NaHCO<sub>3</sub> was double blinded and 129 randomised (block randomisation), as a laboratory technician outside of the research group 130 prepared the NaHCO<sub>3</sub>. Likewise, the time to peak HCO<sub>3</sub><sup>-</sup> was determined by researchers 131 outside of the study and the participant was not informed of their time to peak to ensure the 132 double blind nature of the study. For the PLA condition, a time to peak HCO<sub>3</sub><sup>-</sup> was used from 133 either SBC2 or SBC3.

134

## 135 Four-kilometre cycling protocol, blood measures and perceptual measures

136 The next visit involved a familiarisation to the 4 km cycling TT on a Velotron cycle ergometer 137 (Velotron, RacerMate Inc., USA) interfaced with Velotron coaching software (RacerMate Inc., 138 USA). This ergometer has displayed high test-retest reliability with excellent ICC values of 139 between r = 0.90 to 0.96, p < 0.01 for mean power in TT events (Astorino, 2011; Costa, 140 Guglielmo & Paton, 2017). Participants selected a preferred handlebar and saddle position, 141 whilst they were also permitted to change gears freely throughout each TT using their preferred 142 fixed gear ratios. These settings were then adopted for all subsequent trials. Strong verbal 143 encouragement was provided throughout the TT and feedback on the distance covered and 144 cadence was provided via the software (Stone et al., 2011), but time elapsed was blinded. Time 145 to complete, mean power and mean speed was recorded for both the total distance and 0.5 km 146 splits, along with heart rate (HR) every 0.5 km (Polar, T31, Finland). Blood measures for pH 147 and  $HCO_3^{-1}$  were taken pre-ingestion and post-exercise as per the previously described method. 148 A 5µl sample for blood lactate (BLa) was also taken at the same respective time points (Lactate 149 Pro 2, Arkray, Japan). Ratings of perceived exertion (6-20; Borg, 1982) for the whole body 150 (RPE<sub>0</sub>), legs (RPE<sub>L</sub>), and affective perceptions of work rate (11-point bipolar scale with +5 151 representing 'very good' and -5 representing 'very bad') were recorded every 1 km (Thomas 152 et al., 2015). This procedure was repeated another three times, with the exception that either 0.2 g·kg<sup>-1</sup> BM NaHCO<sub>3</sub> (SBC2), 0.3 g·kg<sup>-1</sup> BM NaHCO<sub>3</sub> (SBC3) or a taste matched placebo 153 154 (PLA) containing 0.07 g.kg<sup>-1</sup> BM sodium chloride (NaCl) was ingested, after baseline measures 155 were taken. Participants then sat quietly rested until their respective predetermined time to peak 156 HCO<sub>3</sub>, at which point a further blood sample was taken. Treatments were administered in a 157 double-blind manner, and for PLA treatments, a time to peak HCO<sub>3</sub><sup>-</sup> time frame from an SBC 158 treatment was selected randomly by a researcher outside of the study to maintain the double-159 blind design. Following ingestion, and up to the individuals respective time to peak HCO<sub>3</sub><sup>-</sup>, GI 160 discomfort was measured using a visual analogue scale (VAS) every 10 min, as per previous 161 studies (Miller et al., 2016; Gough et al., 2017a).

162

## 163 Statistical analysis

164 Assessed variables were analysed using both Shapiro-Wilk tests and standard graphical 165 methods for normality, whilst a Mauchly test was used for homogeneity and 166 variance/sphericity. A paired sampled t-test was used to assess the severity and time to peak 167 GI discomfort between SBC treatments. Both mean power and speed were analysed using a 168 repeated measures ANOVA. Otherwise, a two-way repeated measures ANOVA (e.g. condition 169 x each 0.5 km segment/time point) was used and where either interactions or main effects were 170 observed, Bonferroni corrected posthoc pairwise comparisons were carried out. Where main 171 effects or interactions were observed, partial eta squared  $(P\eta^2)$  effect size is reported. Between 172 treatment effect sizes (d) were calculated using the difference in means divided by the pooled 173 SD of the compared trials (Nagakawa & Cuthill, 2007), however with a Hedge's g bias 174 correction to account for the sample size in this study (Lakens, 2013). All effect size 175 interpretations were considered as trivial (<0.20), small (0.20-0.49), moderate (0.50-0.79) or 176 large ( $\geq 0.80$ ) (Cohen, 1988). Intraclass Correlation Coefficients (ICC) were used to determine 177 the reproducibility of blood metabolites (i.e. time to peak HCO<sub>3</sub><sup>-</sup> and pH) following SBC conditions and are reported with r value and significance value (p value). Interpretation of 178 179 reproducibility was determined by the respective r value with categories of poor (<0.40), fair 180 (0.40-0.59), good (0.60-0.74) and excellent (>0.74). Data are presented as mean  $\pm$  SD with 181 95% confidence intervals (CI) unless otherwise stated. Statistical significance was set at p 182 <0.05 and data were analysed using SPSS v22 for Windows (SPSS Inc., Chicago, IL, USA).

- 183
- 184 **Results**

# 185 *Performance responses for all participants (n =11)*

Faster mean completion times (Figure 1) by  $8.3 \pm 3.4$  s were observed following SBC2 (p < 0.001, CI = 12.0, 4.7, d = 0.64) and by 8.6 s  $\pm 5.2$  s following SBC3 compared to PLA, respectively (p =0.003, CI = 14.2, 3.0, d = 0.66). There was no difference between SBC2 and

- 189 SBC3 (374.0  $\pm$  13.3 vs. 373.7  $\pm$  13.3 s, p =0.87, CI = -3.0, 3.7, d =0.02; Figure 1).
- 190
- 191 \*\*Figure 1 near here\*\*
- 192

193A  $16 \pm 13$  W (+5.7%) increase in mean power was observed following SBC2 ( $304 \pm 28$  W, p194=0.02, CI = 2.6, 30.3, d = 0.62), while in SBC3 an increase of  $16 \pm 15$  W (+5.9%) was observed195( $304 \pm 31$  W, p =0.03, CI = 1.1, 32.9, d = 0.58; Figure 2a) compared to PLA ( $287 \pm 25$  W).196There was no difference between SBC2 and SBC3 (p =0.90, CI = -10.2, 9.1, d = 0.01).197Following SBC2, a  $0.9 \pm 0.6$  km.h<sup>-1</sup> (+2.4%) increase in mean speed was observed compared198to PLA ( $38.6 \pm 1.4$  vs.  $37.7 \pm 1.1$  km.h<sup>-1</sup>, p =0.008, CI = 0.2, 1.6, d = 0.69). Similarly, a  $0.8 \pm 0.6$  km.h<sup>-1</sup> (+2.0%) increase in mean speed was observed following SBC3 ( $38.4 \pm 1.3$ , p =0.02,

200 CI = 0.1, 1.4, d =0.56), whilst there was no difference between SBC conditions (p =0.42, CI = -0.3, 0.6, d =0.14; Figure 2b).

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203 \*\* Figure 2 near here\*\*

204

Performance responses for participants who suffered gastrointestinal (GI) discomfort (n = 8) Despite the occurrence of GI discomfort, SBC2 improved performance by  $9.0 \pm 3.8$  s in SBC2 (p = 0.001, CI = 4.5, 13.5, d = 0.68) and  $8.9 \pm 6.1$  s in SBC3 (p = 0.02, CI = 1.7, 16.2, d = 0.68) compared to PLA. Only one participant failed to improve performance (0.1 s difference vs. PLA), whilst three participants improved by less than the 3 s threshold that was set in the priory power calculation for a meaningful effect (range = 2-2.6 s improvement vs. PLA).

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## 212 Blood metabolite responses

Absolute peak change in  $HCO_3^-$  from baseline was  $5.5 \pm 0.7$  in SBC2 and  $6.5 \pm 1.3$  mmol·l<sup>-1</sup> in SBC3 which was not significantly different (p =0.07; *d* =0.92). Peak  $HCO_3^-$  occurred within a range of between 40 to 110 mins in SBC2 (mean  $62 \pm 20$  min, CV: 33%), and between 40 to 100 min in SBC3 (mean  $73 \pm 20$  min, CV: 27%; Figure 3).

217

218 \*\*Figure 3 near here\*\*

219

The change from baseline to the peak pH was not significantly different between SBC conditions (p =0.13, d =0.75; SBC2 =0.07 ± 0.02, SBC3 =0.09 ± 0.03). In subsequent cycling trials (i.e. 4km TT's) good reproducibility was observed for absolute mean change from baseline in pH following both SBC2 (+0.06; ICC r=0.67, p=0.026) and SBC3 (+0.06; r=0.65, p=0.040). Greater reproducibility was observed for absolute mean change in HCO<sub>3</sub><sup>-</sup> however, displaying excellent reliability in both SBC2 (+4.9 mmol·l<sup>-1</sup>; *r* =0.86, p =0.002) and SBC3
(+5.6 mmol·l<sup>-1</sup>; *r* =0.88, p <0.001).</li>

227

In the cycling trials, a time  $\times$  treatment interaction was observed for pH (p =0.048,  $P\eta^2$ =0.285) 228 229 whereby pH was  $+0.07 \pm 0.02$  (+0.9%) greater at time to peak (figure 4a) for SBC2 ( $7.46 \pm$ 230 0.03; p <0.001, CI = 0.09, 0.04, d = 2.64) and  $0.08 \pm 0.02$  (+1%) greater for SBC3 (7.47 ± 0.02; 231 p < 0.001, CI = 0.09, 0.05, d = 3.85) compared to PLA (7.39 ± 0.02). There was no difference 232 between SBC2 and SBC3 (p = 0.69, CI = -0.3, 0.1; d = 0.38). A time × treatment interaction was observed for HCO<sub>3</sub><sup>-</sup> (p <0.001,  $P\eta^2$  =0.796), with values greater following supplementation of 233 NaHCO<sub>3</sub> (Figure 4b). At time to peak HCO<sub>3</sub><sup>-</sup>, SBC2 was 5.0 mmol·1<sup>-1</sup>  $\pm$  1.0 mmol·1<sup>-1</sup> (+17.6%) 234 235  $(28.6 \pm 1.1 \text{ mmol}^{-1}; \text{ p} < 0.001, \text{ CI} = 6.0, 4.1, d = 5.22)$  and SBC3 was  $5.9 \pm 1.1 \text{ mmol}^{-1}$ (+20.0%) (29.5 ± 1.0 mmol·1<sup>-1</sup>; p <0.001, CI = 6.9, 5.0, d = 6.58) greater than PLA (23.6 ± 0.7) 236 237 mmol·1<sup>-1</sup>). There was no difference between SBC2 and SBC3 (p = 0.34, CI = -2.3, 0.6, d = 0.82). 238

239 Post exercise HCO<sub>3</sub><sup>-</sup> was  $\pm 1.8 \pm 1.3$  mmol·l<sup>-1</sup> ( $\pm 12.3\%$ ) greater for SBC2 ( $16.0 \pm 2.2$  mmol·l<sup>-1</sup>; 240 p =0.004, CI = 2.9, 0.6, d =0.79), and +1.5 ± 1.3 mmol·1<sup>-1</sup> (+10.9%) greater for SBC3 (15.8 ± 2.7 mmol·1<sup>-1</sup>; p =0.01, CI = 2.7, 0.4, d = 0.62) compared to PLA (14.2 ± 2.2 mmol·1<sup>-1</sup>). There 241 was a main effect for treatment in HCO<sub>3</sub><sup>-</sup> change during exercise (p <0.001,  $P\eta^2 = 0.714$ ), 242 243 whereby the change in HCO<sub>3</sub><sup>-</sup> was  $3.3 \pm 1.8$  mmol<sup>1-1</sup> (+25.9%) greater following SBC2 (12.7 244  $\pm 2.6 \text{ mmol}^{1-1}$ ; p =0.001, CI = 4.9, 1.6, d =1.37) and 4.4  $\pm 1.7 \text{ mmol}^{1-1}$  (+31.7%) greater for 245 SBC3 (13.8 ± 2.7 mmol<sup>-1</sup>; p <0.001, CI = 5.9, 2.8, d =1.78) compared to PLA (9.4 ± 2.0 mmol·1<sup>-1</sup>). There was no difference between SBC conditions (p =0.59, CI = -1.2, 3.3; d =0.40). 246 A main effect for time was observed for BLa (p <0.001,  $P\eta^2 = 0.957$ ) with all conditions 247 248 displaying greater post-exercise BLa compared to pre-exercise (Figure 4c). Post-exercise, a time × treatment interaction was observed for BLa (p <0.001,  $P\eta^2 = 0.577$ ) as SBC2 was +3.7 249

250  $\pm 2.8 \text{ mmol}\cdot l^{-1}$  (+22.5%) greater than PLA (16.1  $\pm 3.4 \text{ vs.} 12.5 \pm 2.7 \text{ mmol}\cdot l^{-1}$ , p =0.006, CI = 251 1.1, 5.8, *d* =1.13; Figure 4c), with SBC3 greater by +3.7  $\pm 2.4 \text{ mmol}\cdot l^{-1}$  (+22.7%) (16.1  $\pm 3.4$ 252 mmol}\cdot l^{-1}; p =0.002, CI = 1.5, 5.8, *d* =1.13). No differences between SBC conditions were 253 evident for post-exercise BLa (p =0.61, CI = -2.3, 2.2; *d* =0.01).

254

- 255 \*Figure 4 near here\*\*
- 256
- 257 Gastrointestinal (GI) discomfort

Four participants reported symptoms of belching and stomach bloating in SBC2, compared to seven participants reporting symptoms of belching, stomach cramp, bowel urgency and diarrhoea in SBC3. There was no significant difference in severity of GI discomfort between SBC treatments (SBC2 =1.4 ± 1.5 vs. SBC3 =4.6 ± 3.6; p =0.10), although a large effect size was evident (d = 0.88). Similarly, time to peak GI discomfort was not significantly different between SBC treatments (SBC2 =20 ± 24 vs. SBC =43 ± 31min, p =0.13), although revealed a large effect size (d = 0.80).

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# Heart rate (HR), ratings of perceived exertion (RPE) and affective perceptions of work rate scale

Heart rate was unaffected by NaHCO<sub>3</sub> ingestion as no time × treatment interaction was observed (p =0.56,  $P\eta^2$  =0.055). There was a main effect for time (p <0.001,  $P\eta^2$  =0.977) for HR and mean data combined from all treatments displayed HR at 500m was 144 ± 3 b min<sup>-1</sup>, compared to 171 ± 2 b min<sup>-1</sup> at 4 km, respectively. A main effect for time was observed for RPE<sub>0</sub> (p <0.001,  $P\eta^2$  =0.849), as at 1 km RPE<sub>0</sub> was 14 ± 1 compared to 17 ± 1 at 4 km, although no time × treatment was apparent (p =0.31,  $P\eta^2$  =0.109). A main effect for time was observed for RPE<sub>L</sub> (p <0.001,  $P\eta^2$  =0.657), as at 1 km RPE<sub>L</sub> was 15 ± 1 compared to 18 ± 0 at 4 km, although no time × treatment interaction was evident (p =0.73,  $P\eta^2$  =0.085). Affective perceptions of work rate revealed no time × treatment interaction (p =0.38,  $P\eta^2$  =0.099) or main effect for time (p =0.92,  $P\eta^2$  =0.020).

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#### 279 Discussion

In agreement with our hypothesis, this study reports that both 0.2 gkg<sup>-1</sup> (SBC2) and 0.3 gkg<sup>-1</sup> 280 281 BM (SBC3) NaHCO<sub>3</sub> improves 4 km TT cycling performance in trained cyclists when 282 individualised to a predetermined time to peak  $HCO_3^-$ . Time to complete the time trial was 283 2.2% faster in SBC2 and 2.3% in SBC3 compared to PLA, whilst there was also no statistical 284 difference between SBC conditions suggesting both amounts are appropriate to enhance this 285 type of exercise performance. Combining such performance effects with the reduced instances and severity of GI discomfort following 0.2 gkg<sup>-1</sup> BM NaHCO<sub>3</sub> however, the present study 286 287 findings suggest this amount may be more attractive to the athlete in a practical setting.

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289 The findings of the present study contrast that of two recent studies reporting no effect of 290 NaHCO<sub>3</sub> on 4 km TT performance (Callahan et al., 2017; Correia-Oliveira et al., 2017). Indeed, 291 Callahan et al. (2017) reported a 'possibly trivial' effect and Correia-Oliveira (2017) reported no significant supplement interaction in ANOVA analysis following 0.3 gkg<sup>-1</sup> BM NaHCO<sub>3</sub>. 292 293 In comparison, the present study displayed a statistically significant effect and a moderate 294 effect size for both SBC2 and SBC3. This ergogenic effect was most likely realised due to 295 supplementing NaHCO<sub>3</sub> to a predetermined time to peak  $HCO_3^{-1}$ , as this would have ensured 296 peak bioavailability of HCO<sub>3</sub><sup>-</sup> at the commencement of exercise. In particular, the increase in 297 HCO<sub>3</sub><sup>-</sup> following the SBC2 treatment of the present study was similar, whilst the SBC3 treatment was superior, to the values reported in the aforementioned studies with 0.3 g kg<sup>-1</sup> BM 298 NaHCO<sub>3</sub> (SBC2 = 4.9 to 5.5 mmol<sup>-1</sup>, SBC3 = 5.6 to 6.5 mmol<sup>-1</sup> vs. Callaghan et al. = +3299

300 mmol·l<sup>-1</sup> vs. Correia-Oliveira et al. = +5mmol·l<sup>-1</sup>). Based on this evidence, it is therefore more 301 appropriate to identify time to peak HCO<sub>3</sub><sup>-</sup> prior to the use in exercise to elicit ergogenic effects 302 on performance. A consideration, however, is that identifying time to peak HCO<sub>3</sub><sup>-</sup> presents a 303 logistical challenge, as this would require a visit to a laboratory or access to a portable blood 304 gas analyser.

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306 A unique finding of the present study was the lack of a dose-dependent effect on exercise 307 performance, with SBC3 improving performance to a similar magnitude as SBC2. These findings are in contrast to McNaughton (1992), reporting 0.3 g·kg<sup>-1</sup> BM NaHCO<sub>3</sub> improved 308 309 TWD greater than 0.2 g·kg<sup>-1</sup> BM NaHCO<sub>3</sub> during 60 seconds of maximal cycling compared to 310 a placebo. The negligible 0.1% difference observed between SBC2 and SBC3 are more in 311 agreement with the findings of McKenzie et al. (1986) reporting a 0.3% difference between 312 0.15 gkg<sup>-1</sup> BM and 0.3 gkg<sup>-1</sup> BM NaHCO<sub>3</sub>. Individual performance responses did reveal that 313 three participants improved to a greater extent in SBC2 compared to SBC3, whilst two 314 participants improved to a greater extent in SBC3 compared to SBC2 based on the 3 s cut off 315 from the prior power calculation. These data combined suggest lower amounts of NaHCO<sub>3</sub> (i.e. 0.2 g·kg<sup>-1</sup> BM) are likely to be sufficient to enhance exercise of this duration and intensity, 316 317 although athletes should trial each dose prior to use in competition to evaluate which amount 318 of NaHCO<sub>3</sub> provides a larger ergogenic benefit. Likewise, considering the potential for the 319 onset of GI discomfort, athletes who are susceptible to such symptoms should conduct a 320 risk:benefit analysis of NaHCO<sub>3</sub> supplementation.

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322 It is purported that mitigating the severity of GI discomfort is important to obtain a performance
 323 benefit following NaHCO<sub>3</sub> supplementation, as Saunders et al. (2014) reported a significant
 324 effect on performance only upon the removal of participants who suffered from GI discomfort.

325 The present study findings contrast this by reporting a significant 2.3% improvement following 326 both SBC2 and SBC3, despite the occurrence of mild to moderate GI discomfort. Reasons for 327 this may be due to the good tolerance of NaHCO<sub>3</sub> in our participant cohort, although it is difficult to compare with the work of Saunders et al. (2014) as no explicit statistical analysis 328 329 on GI discomfort is available. Nonetheless, there may still be a relationship between GI 330 discomfort and performance, as for instance, participant 8 in the present study suffered from 331 moderate diarrhoea and bowel urgency in SBC3 and no improvement in performance was 332 observed (0.1 s). While performance in SBC2 was improved by 8.9 s in the same participant 333 when no instances of GI discomfort occurred. Combining this finding with other investigations 334 where participants have self-withdrawn, or have been withdrawn by the research team due to 335 the severity of GI discomfort, the responses from NaHCO<sub>3</sub> still warrant observation in training 336 prior to use in competition (Gough et al., 2017a, 2017b; Jones et al., 2016). Nonetheless, 337 smaller amounts of NaHCO<sub>3</sub> may be an attractive solution to the athlete to reduce the severity 338 of GI discomfort symptoms whilst still providing ergogenic effects to exercise performance.

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340 The enhancements of acid base balance following NaHCO<sub>3</sub> are the most likely mechanism for 341 an improved performance in the present study, as both SBC2 and SBC3 raised HCO<sub>3</sub><sup>-</sup> and pH 342 significantly compared PLA. An increase in extracellular HCO<sub>3</sub><sup>-</sup> is suggested to increase H<sup>+</sup> 343 efflux during exercise due to the up-regulation of the lactate/H<sup>+</sup> cotransporter, leading to 344 increased provision of anaerobic energy contribution (Marx et al., 2002). The change in HCO<sub>3</sub>-345 was superior in both SBC2 (+25.9% vs. PLA) and SBC3 (+31.7% vs. PLA) whilst post-346 exercise blood lactate was also significantly higher (~15%) in the SBC conditions. These 347 changes in blood acid base balance and BLa are indicative of exercise at higher exercise 348 intensities in the SBC conditions and hence, improved performance. Furthermore, between 349 SBC conditions there were minimal differences in respect of blood metabolites changes prior to, or during exercise. This provides an explanation why there were no dose-dependent effectson performance in the present study.

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# 353 Conclusion

354 Ingestion of NaHCO<sub>3</sub> individualised to time to peak HCO<sub>3</sub><sup>-</sup> improves 4 km TT cycling performance in trained cyclists. Ingestion of both 0.2 g kg<sup>-1</sup> BM and 0.3 g kg<sup>-1</sup> BM NaHCO<sub>3</sub> 355 356 equally increase buffering capacity and subsequently provided ergogenic benefits to exercise 357 performance. No difference was observed between SBC conditions; therefore, athletes can plausibly use a lower amount of NaHCO<sub>3</sub> (i.e. 0.2 g·kg<sup>-1</sup> BM) particularly if they are susceptible 358 359 to the onset GI discomfort. Future research should investigate the dose-dependent effects of both 0.2 gkg<sup>-1</sup> BM and 0.3 gkg<sup>-1</sup> BM NaHCO<sub>3</sub> during exercise of different intensities and 360 361 durations.

#### **Disclosure statement**

The authors report no conflicts of interest.

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# List of figures

Figure 1 – Mean ( $\pm$ SD), and individual 4 km time trial performance times following each condition. \*denotes significantly different from PLA (p <0.05).

Figure 2 – Mean ( $\pm$ SD) cycling power (A) and speed (B) during each 0.5 km segment of the time trial. Significant increase (p <0.05) in SBC2 = # and SBC3 = ## compared to PLA.

Figure 3 – Individual time to peak blood bicarbonate (HCO<sub>3</sub><sup>-</sup>) following SBC2 and SBC3.

Figure 4 – Mean ( $\pm$ SD) blood pH (A), bicarbonate (HCO<sub>3</sub><sup>-</sup>) (B) and lactate (C) responses during experimental treatments. Significantly different (p <0.05) in SBC2 = # and SBC3 = \* compared to PLA.