Accepted Manuscript

Type of manuscript: Regular Article

Title: Carbohydrate mouth rinse during physical activity to improve cognitive function:

A randomized cross-over trial

Author(s): Donghyun KIM *¹* *, Tsuyoshi WADAZUMI*²*

Affiliation(s):

1 Graduate School of Health and Well-being, Kansai University, Osaka, Japan, Email: kdh3970@gmail.com *²*Faculty of Health and Well-being, Kansai University, Osaka, Japan, Email: wadazumi@kansai-u.ac.jp

Running title: Carbohydrate mouth rinse for cognitive function

Number of Figures: 4

Number of Tables: 3

***Corresponding author:** Donghyun KIM, Graduate School of Health and Well-being, Kansai University, Osaka, Japan **Phone:** +81-(0)70-4412-8441 **Email:** kdh3970@gmail.com

Abstract

The dorsolateral prefrontal cortex (DLPFC) is a brain region responsible for executive function. Recent studies have reported that low-intensity physical activity (LPA) can activate this region and improve executive function. Carbohydrate mouth rinse (CMR), an ergogenic conditioning method in sports science, has been shown to activate the DLPFC. This study hypothesized that LPA-induced improved executive function could be synergistically enhanced by CMR. We investigated the impact of combining LPA with CMR to improve executive function.

Seventeen healthy university students (eight males, nine females) participated. For LPA, instead of commonly-used exercise modalities, such as bicycle ergometers or treadmills, we employed a seated dual-task exercise (DE) that involved performing cognitive tasks and physical activity simultaneously, which is considered effective for cognitive function. Four trials were conducted using a crossover design: trial 1 control (CON), trial 2 (CON+CMR), trial 3 (DE), and trial 4 (DE+CMR). The Trail Making Test Type B (TMT-B, Japanese version) and Stroop Test (ST, Japanese version) assessed cognitive function, and CMR was administered with a 6% glucose solution four times during CON and DE.

The results indicated a significant interaction in Interference Rate I of the ST, showing improved performance due to DE. An interaction was also observed in Interference Rate II, highlighting enhanced performance when DE was combined with CMR. Although no interaction was found in the TMT-B, a pre-post comparison revealed improved performance in the DE + CMR trial.

These findings suggest the potential for further improvement in executive function when DE is combined with CMR.

Keywords: carbohydrate, sedentary behavior, dual-task exercise, cognitive function

論文名

デュアルタスク運動とマウスリンスの併用が認知機能に及ぼす影響(ランダム 化比較試験: クロスオーバーデザイン)

著者

 \Leftrightarrow 東鉉¹、弘原海 剛²

¹関西大学大学院 人間健康研究科、大阪、日本

²関西大学 人間健康学部、大阪、日本

抄録

背外側前頭前野(DLPFC)は実行機能を司る脳領域であり、近年では低強度 身体活動(LPA)でも DLPFC が活性化し、実行機能向上に効果的であることが報告 されている。一方、炭水化物マウスリンス(CMR)は、近年スポーツ科学分野 で用いられるようになったエルゴジェニックなコンディショニング法であり、 口に炭水化物水溶液を含むだけで、DLPFC を活性化することが明らかにされて いる。そこで、本研究では LPA に CMR を介入すれば更なる実行機能向上が見ら れると仮説を立て、LPA に CMR を組み合わせることで実行機能の向上に及ぼす 影響について明らかにすることを目的とした。対象は健康な大学生17人(男性 8 人、女性 9 人)。LPA としては一般的に用いられる自転車エルゴメーターやト

レッドミルではなく、運動と認知課題を同時に行うことで認知機能向上に効果 的と言われている座位姿勢デュアルタスク運動(DE)を採用し、以下の 4 試行 をクロスオーバー法で実施した(試行1コントロール(CON)、試行2 CON+CMR、 試行 3 DE、試行 4 DE+CMR)。認知機能検査には日本語版の Stroop Test (ST) と Trail Making Test Type B(TMT-B)を用いた。CMR には 6%グルコース水溶 液を用いて CON と DE 試行中に全 4 回実施した。ST の干渉率Ⅰでは交互作用が 認められ、DE による成績向上が認められた。また、干渉率Ⅱでは交互作用が認 められ(p<0.010)、DE と CMR を組み合わせるによる成績向上が認められた。ま た、TMT-B では交互作用が認められなかったが、前後比較では DE+CMR 試行で成 績向上が認められた (p<0.001)。

これらのことから、DE中にCMRを組み合わせることで更なる実行機能向上の 可能性が示唆された。

Introduction

 The 2020 WHO guidelines state that any form of physical activity (PA) is beneficial for health even when performed for short durations. Additionally, owing to the adverse health effects of prolonged sitting, it is recommended to perform PA of any intensity, 5 including light-intensity.¹⁾ Performing regular PA has been shown to improve physical health, alleviate symptoms of depression and anxiety, as well as enhance mental health 7 and cognitive function.^{1,2)}

 The effects of acute PA interventions on cognitive function have been reported in 9 several previous studies.³⁻⁷⁾ As an intervention mechanism, PA has been shown to activate the primary motor cortex, supplementary motor area (SMA), parietal lobe (PL), and areas of the frontal lobe, including the dorsolateral prefrontal cortex (DLPFC)—one of the 12 brain regions involved in cognitive activities.^{3,4,8)} Specifically, the DLPFC is responsible 13 for executive functions.⁹⁾ Executive functions refer to the processes necessary for the planning, action, and thinking required to achieve goals. Elements that constitute 15 executive functions include inhibition, working memory, and cognitive flexibility.¹⁰⁾ Therefore, since the DLPFC is crucial for performing executive functions, its activation is key to improving cognitive function.

 Students today spend prolonged periods sitting in classrooms, necessitating strategies 19 such as replacing this time with PA .¹¹⁾ Programs such as Physical Activity Across the Curriculum (PAAC) and Take10! have been developed and implemented as optimal PA interventions in the classroom, resulting in reported educational benefits and improved 22 cognitive function.^{12,13)} However, concerns have been raised about preparing an 23 appropriate space for these activities in the classroom.¹¹⁾

One easily accessible form of light-intensity physical activity (LPA) is dual-task

 exercise (DE), which involves performing two tasks simultaneously, such as cognitive tasks and LPA. Since both LPA and cognitive tasks induce brain activation, combining 27 them enhances cognitive function.¹⁴⁾ Studies targeting older adults have reported 28 numerous benefits of DE on cognitive function.¹⁵⁻¹⁷⁾ However, research on the effects of 29 DE on cognitive function in young people is limited.¹⁸⁻²⁰⁾

 Carbohydrate mouth rinse (CMR) has emerged as an ergogenic conditioning method used in sports science. Ingesting food during exercise for energy replenishment can cause gastrointestinal discomfort and stomach pain. However, CMR involves rinsing the mouth with a carbohydrate solution without ingestion, thus eliminating these concerns. Numerous previous studies in sports science have demonstrated the effectiveness of CMR 35 in improving exercise performance, $21-24$ particularly during high-intensity intermittent exercise. CMR works by central fatigue inhibition rather than maximal strength enhancement, and has been proven effective for endurance activities. As an intervention mechanism, the effects of CMR on different brain regions (DLPFC, ventral striatum, anterior cingulate cortex [ACC], orbitofrontal cortex [OFC]) have been mediated 40 centrally.²¹⁾

 Recent research has focused on the activation of the DLPFC by CMR intervention during exercise and its impact on cognitive function. However, these previous studies applied moderate to high exercise intensity, while none used light-intensity exercise.

44 Since the effects of CMR on brain activity occur even without fatigue, we hypothesized that CMR intervention during LPA would synergistically lead to further improvement in executive function. Therefore, this study aimed to investigate the effects of CMR intervention during DE on executive function in young adults using short-duration LPA performed in a seated position, which is an activity easily accessible to

anyone.

Materials and methods

Participants

 The sample size was calculated using G*Power 3.1.9.4 (Institute for Digital Research and Education, Düsseldorf, Germany). Utilizing an F test for repeated measures with within-between interactions, an effect size of 0.25, a significance level of 5%, and a power of 80%, the a priori power analysis indicated that the required sample size was 12 participants per group.

 The participants included 17 healthy university students (8 males and 9 females) who were recruited from July 23 to 30, 2021. Their physical characteristics are presented in Table 1.

(Table 1)

Block exercise

 Block exercise (BE; "Co-kara"), Sakai City's version of dementia prevention 63 exercises, was used for $DE^{.15)} BE$ is a low-intensity exercise performed in a seated 64 position, with an intensity of $2-3$ METs.²⁶⁾ BE consists of hand and leg movements (blocks) synchronized to music (Figure 1). The exercise allows for various movement patterns and rhythms, and can be performed to any preferred music, which helps maintain engagement and prevents boredom. One component of music is rhythm, and rhythmic activities, such as finger tapping, have been reported to activate brain regions including 69 the DLPFC.²⁷⁻²⁹⁾ Based on this evidence, our study employed BE, an exercise synchronized with rhythm, to investigate its effects. In this study, heart rate was measured 71 during BE, and exercise intensity was calculated using the Karvonen method. The results showed a range of 3.9-21.2% HRmax, which was confirmed to meet the criteria 73 for low-intensity exercise (\leq 57% HRmax) as suggested by the ACSM ³¹⁾.

(Figure 1)

Cognitive function task

 Although various testing methods exist for evaluating cognitive function, we used the Japanese version of the New Stroop Test Ⅱ (ST) and the Trail Making Test Type B (TMT-B), commonly used to measure executive function.

 The ST evaluates an individual's ability to suppress interference from two pieces of information—letter meaning and letter color—and to make an attentional choice. The interference includes Stroop interference (SI), which eliminates letter meaning and reads 82 letter color, and reverse-Stroop interference (RI), which eliminates letter color and reads letter meaning.

 The New ST-II³²⁾ consists of four tasks: reverse Stroop control condition, reverse SI condition, Stroop control condition, and SI condition. The control condition matches letter color and meaning, and the interference condition mismatches letter color and meaning. Each task consists of 10 practice tasks and 100 main tasks. The main task is performed for one minute. The interference rate is calculated from the results, which confirm the Stroop effect that occurs in Tasks 2 and 4 (SI conditions: incongruent) compared to Tasks 1 and 3 (control conditions: congruent). The formula for calculating the interference rate is shown below (Figure 2-B).

- RI rate (Interference Rate I):
- [(number of correct answers for Task 1 number of correct answers for Task 2)/
- number of correct answers for Task 1] x 100
- SI rate (Interference Rate II):
- [(number of correct answers for Task 3 number of correct answers for Task 4)/

number of correct answers for Task 3] x 100

 The TMT consists of two types of tests: TMT-A and TMT-B. TMT-A tests an individual's ability to connect numbers from 1 to 25 in sequence and mainly requires visual perception ability. The TMT-B primarily assesses working memory but also 101 assesses task-set switching ability³³⁾ (Figure 2-A). The Japanese version of the TMT-B was used in this study.

 Participants performed the cognitive function test on a Microsoft Surface Go 3 tablet by touching the screen. In addition, the participants practiced the TMT-B and ST several times beforehand.

(Figure 2)

Experimental protocol

 The experiment was conducted simultaneously for each participant, and each trial was separated by an interval of at least three days. The participants were instructed not to engage in strenuous exercise or drink alcohol on the day before the experiment and not to consume anything other than water four hours before the start of the experiment. Once they arrived at the laboratory on the day of the experiment, the participants were instructed to rest in a seated position for 30 minutes in the center of a space surrounded by partitions on all four sides. They were asked to maintain a comfortable posture against the backrest of the chair until the end of the experiment. Heart rate was also measured using a polygraph system (RMT-1000, Nihon Kohden) and LabChart pro software (LabChart 8, ADInstruments) to evaluate exercise intensity.

118 The experiment consisted of four trials: Trial 1 [control (CON)], Trial 2 (CON+CMR),

Trial 3 (DE), and Trial 4 (DE+CMR). The CON and CON+CMR procedures consisted of

resting (five minutes) followed by performing three (TMT-B) tests and four different STs

 (see Figure 3). While in the sedentary posture, the participants then listened quietly to three music pieces, each about four minutes long. They rested for two minutes and completed the same cognitive function tests. DE and DE+CMR followed the same protocol as CON, but DE was performed with participants watching a recorded video during the experiment. Three songs (approximately 12 minutes in total) were chosen as the exercise duration because previous studies have reported that 11 to 20 minutes of 127 exercise improves cognitive function regardless of the exercise intensity.³⁴⁾ A medium speed tempo (moderato) was selected for the three songs: 92, 101, and 106 beats per minute, respectively.

 We conducted a crossover randomized controlled experiment (1-2-3-4, 2-3-4-1, 3-4- 1-2, 4-1-2-3) with four trials divided into four groups of participants. A sufficient number of test trials were conducted before the experiment to familiarize the participants with the experimental procedures, cognitive functions, and DE.

(Figure 3)

Carbohydrate mouth rinse

 Although the carbohydrate solution was placed in the mouth and spat out in previous 137 CMR studies, recent studies have proven that the spray method is equally effective.²⁴⁾ A 6% concentration carbohydrate solution was used in this study based on previous 139 research.³⁵⁾ The CMR solution, which consisted of 30 g of glucose (manufactured by FUJIFILM Wako Pure Chemical Corporation) dissolved in 500 ml of water, was placed in a spray container (13 ml). The point of CMR implementation was indicated by arrows in Trials 2 and 4 (Figure 3). The participants were instructed to administer six pumps of solution into their mouth using the spray container, and not to swallow (Figure 4). (Figure 4)

Analytic methods

146 The TMT-B was conducted three times each in both the pre- and post-treatment tasks. To reduce measurement error, the intraclass correlation coefficient (3,1) was used to detect the reliability of the three trials and the top two trials. Although reliability was over 0.5 in the top two trials, reliability over 0.4 was not detected in all three trials; therefore, the average of the top two scores was used for analysis (0.4 slight, 0.41–0.6 fair, 0.61–0.8 moderate, 0.81–1.00 almost perfect). The ST was analyzed using Interference Rates I and II.

 The TMT-B performance time and ST Interference Rates I and II for each trial were compared by calculating the amount of change between the pre- and post-intervention results.

Statistical processing

 SPSS Statistics 27 (IBM Corp., Armonk, NY, USA) was used for statistical processing. The data collected were verified for normality using the Shapiro–Wilk test 159 and presented as mean \pm standard deviation. Comparisons between trials were conducted 160 using a two-way repeated measures ANOVA (Trial \times Time). When interaction or main effects were observed, simple main effect tests and multiple comparison tests were used for further analysis. Effect sizes for ANOVA were analyzed using eta squared values (0.01 < small, 0.06 < medium, 0.14 < large). Bonferroni correction was applied for multiple comparisons while for outcomes where the ANOVA did not reveal a significant interaction effect, exploratory paired t-tests were conducted to examine pre-post 166 differences within each condition. Effect sizes were analyzed using Cohen's d values (0.2) < small, 0.5 < medium, 0.8 < large). Statistical significance was set at a 5% significance level.

Ethical considerations

 The participants were fully informed of the study's purpose and methods. Once they understood these aspects, we obtained their written and verbal consent to participate in the experiment voluntarily. This study was conducted with the approval of the Ethics Committee of the Faculty of Health and Well-being, Kansai University (Ethics Review Number: 2021-08) and registered at the University Hospital Medical Information Network Center (UMIN Center) (UMIN Study No.: UMIN000054101).

Results

(Table 2) (Table 3)

 Cognitive function tests were conducted using ST and TMT-B. A two-way repeated 179 measures ANOVA (Trial \times Time) for Interference Rate I in ST revealed an interaction 180 effect (F = 4.719, p = 0.006, η^2 = 0.228). Additionally, main effects were found for both 181 Trial (F = 6.739, p = 0.001, η^2 = 0.296) and Time (F = 5.267, p = 0.036, η^2 = 0.248). Subsequent post hoc analysis, including calculation of pre- and post-Interference Rate I changes and multiple comparisons, revealed significant differences between CON and 184 DE ($p = 0.027$) as well as between CON and DE+CMR ($p = 0.024$) (Table 2). Paired t- tests of pre- and post-performance revealed significant cognitive improvements in both 186 DE (t $[16] = 2.296$, $p = 0.036$, $d = 0.557$) and DE+CMR (t $[16] = 3.171$, $p = 0.006$, $d =$ 0.769) trials (Table 3).

188 For Interference Rate II, the two-way repeated measures ANOVA (Trial \times Time) also 189 showed an interaction effect (F = 4.246, p = 0.010, η ² = 0.210). While no main effect was 190 found for Trial (F = 1.977, p = 0.130, η ² = 0.110), a main effect was observed for Time (F 191 = 8.929, $p = 0.009$, $\eta^2 = 0.358$). Post hoc analysis on changes in pre- and post-Interference 192 Rate II values and multiple comparisons indicated a significant difference between CON 193 and DE+CMR $(p = 0.041)$ (Table 2). Paired t-tests of pre- and post-performance 194 established significant cognitive improvements in both DE (t $[16] = 2.413$, p = 0.028, d = 195 0.585) and DE+CMR (t $[16] = 5.135$, $p < 0.001$, $d = 1.245$) trials (Table 3).

196 For TMT performance, a two-way repeated measures ANOVA (Trial \times Time) showed 197 no interaction effect (F = 2.249, p = 0.095, η^2 = 0.123) (Table 2); a main effect for Time 198 was observed (F = 13.865, p = 0.002, η ² = 0.464). Exploratory paired t-tests revealed a 199 significant difference between pre- and post-performance in the DE+CMR trial (t $[16] =$ 200 4.961 , $p < 0.001$, $d = 1.189$) (Table 3).

201 **Discussion**

202 We aimed to clarify the impact of combining DE, a seated form of LPA that is easy 203 for anyone to engage in, with CMR on executive function in young adults.

204 The results showed that DE alone improved executive function, and the addition of 205 CMR during DE suggested the potential for further enhancement in executive function.

206 *Effects of DE*

 The effects of acute PA interventions on cognitive function improvement have been reported in numerous studies employing various types and intensities of exercise conditions. The mechanisms underlying these effects include the activation of the DLPFC, increased cerebral blood flow, and the secretion of neurotransmitters, such as 211 catecholamines^{36,37)} and brain-derived neurotrophic factor (BDNF).³⁷⁾ Prior research using functional near-infrared spectroscopy has reported enhancements in both DLPFC 213 activation and cognitive functions following PA interventions.^{3,4,6)} In a previous study, the activation of the DLPFC during DE was measured using near-infrared spectroscopy, and 215 this activation was also reported.¹⁵⁾ This brain activation is likely due to increased

 metabolic activity in the brain regions stimulated by exercise, which has been shown to 217 positively impact cognitive functions.)

 Inhibition, an executive function necessary for performing the ST, has been shown to 219 improve with physical activity (PA) in young adults.³⁹⁾ This suggests that the improvement in ST performance observed in this study can be attributed to this effect. The ST employed in this study activates brain regions such as the DLPFC, SMA, and PL during tasks associated with Interference Rate I. Conversely, the task characteristics for 223 Interference Rate II primarily report DLPFC activation.³²⁾ In the multiple comparisons of the ST, significant improvements were found between the CON and DE groups for 225 Interference Rate I ($p < 0.027$), while no significant improvement was observed for Interference Rate II (Table 2). This may be due to differences in task characteristics. The overlapping activation of brain regions, including the DLPFC, SMA, and PL, during 228 Interference Rate I,³²⁾ suggests a strong influence of exercise on these areas.^{3,4,8)} However, since only the DLPFC is primarily activated during the tasks associated with Interference 230 Rate II ,³²⁾ it is inferred that there were fewer regions activated by exercise compared to Interference Rate I.

 Working memory, which is a necessary executive function for performing the TMT-233 B, has been shown to improve in young adults through $PA.^{39}$. However, the results obtained from TMT-B in this study did not show significant interaction effects and main effects of trial conditions. This lack of significant findings may be due to the favorable results of TMT-B. Previous studies have reported that changes in cognitive function due 237 to PA interventions are less likely to manifest when the pre-test scores are already high.⁴⁰⁾ In this study, the participants' pre-test scores were already higher than the typical TMT-B 239 scores for their age group,⁴¹⁾ which likely explains the absence of significant cognitive improvements between trials. This high performance may be attributed to the fact that participants underwent sufficient test trials before the experiment to familiarize themselves with the procedures and cognitive assessments. On the other hand, the intervention effect on TMT could not be confirmed from the results of the current experiment, but the main effect of time was significant, and the effect size in the DE+CMR condition was larger than in the other conditions, thus suggesting the possibility of a synergistic effect (Table 3).

247 DE has been reported to be effective in improving cognitive function.^{18,19,42,43)} However, to date, most studies on DE have included cognitive tasks during biking or walking. Therefore, the present study is significant as it reveals the effects of DE combined with music on improving cognitive function.

Effects of CMR

 The brain is activated by CMR when the taste buds sense carbohydrates, and information is transmitted through the ACC to the DLPFC and OFC, which activate these 254 areas.⁴⁴⁾ CMR alone did not lead to brain activity indicating improved cognitive function, 255 and no effect on improvement in cognitive function was observed.⁴⁵⁾ This was consistent with the results of the CON+CMR trial in this study. In previous studies that implemented a CMR intervention during moderate to high intensity or prolonged exercise, 258 improvement in cognitive function and suppression of decline were demonstrated.^{46,47)} However, as no studies have previously examined the effects of a CMR intervention during short-duration LPA, this study investigated the topic using DE.

 As a result, in the multiple comparisons of the ST, significant differences were observed between the CON and DE groups, as well as between the CON and DE+CMR groups, for Interference Rate I. However, no significant difference was found between

 DE and DE+CMR, indicating that the effects of exercise were strong enough to obscure the distinction of CMR's impact. By contrast, for Interference Rate II, no significant improvement was noted between the CON and DE groups, but a significant improvement 267 was observed between the CON and DE+CMR groups $(p < 0.041)$ (Table 2). This suggests that the addition of CMR to DE may have led to higher activation in brain 269 regions, such as the DLPFC, ACC, and OFC, which are activated during the ST^{32} and by CMR ^{21,32)} Particularly, the ACC showed strong activation during tasks involving 271 interference⁴⁸⁾; the activation of the ACC and DLPFC due to CMR may be significantly pronounced during the ST.

 Additionally, no improvement in executive function was observed with either CMR or DE alone in the TMT-B. However, the effect size of the DE+CMR condition was greater than that of the other conditions, suggesting the possibility of a synergistic effect using DE and CMR (Table 3). This improvement may be attributed to the priming effect 277 of CMR.⁴⁹⁾ The priming effect refers to how preceding stimuli can facilitate or inhibit subsequent stimuli. This study hypothesized that the preceding stimulus, DE, enhanced the effects of the subsequent stimulus, CMR.

 Based on these findings, it is suggested that combining CMR with low-intensity, short-duration DE may lead to further improvements in executive function, without the necessity for moderate to high-intensity or prolonged exercise. However, this hypothesis remains speculative and requires validation based on physiological data, hence conducting further research to systematically examine these assumptions is crucial.

Limitations

 This study has some limitations. The activation of the DLPFC was not directly measured, preventing a direct comparison with the mechanisms demonstrated in previous

 research. Therefore, further studies are needed to confirm the mechanisms suggesting the involvement of the DLPFC. Additionally, the absence of a CMR placebo solution trial is noted as another limitation, which should be considered in future research.

Perspective

 The CMR used in this study can be applied in various educational settings and to improve the cognitive function of older adults and people with disabilities because this intervention is easily implemented during PA. In addition, DE performed in a seated position is easy for most people, and can be applied universally for those with disabilities or movement limitations. Future studies on CMR interventions during long-term PA and its effects on older adults are necessary.

Conclusions

 The combination of short-duration DE and CMR in young adults suggests the potential for further improvements in executive function.

Acknowledgments

 The authors would like to thank the Kansai University students for participating in the experiment.

References

1. Bull FC, Al-Ansari SS, Biddle S, Borodulin K, Buman MP, Cardon G, Carty C, Chap

ut JP, Chastin S, Chou R, Dempsey PC, DiPietro L, Ekelund U, Firth J, Friedenreich CM

, Garcia L, Gichu M, Jago R, Katzmarzyk PT, Lambert E, et al. 2020. World Health Org

anization 2020 guidelines on physical activity and sedentary behaviour. Br J Sports Med

54(24):1451-62. doi: 10.1136/bjsports-2020-102955.

2. Donnelly JE, Hillman CH, Castelli D, Etnier JL, Lee S, Tomporowski P, Lambourne

K, Szabo-Reed AN. 2016. Physical activity, fitness, cognitive function, and academic ac

 hievement in children: a systematic review. Med Sci Sports Exerc 48(6):1197-222. doi: 10.1249/mss.0000000000000901.

- 3. Yanagisawa H, Dan I, Tsuzuki D, Kato M, Okamoto M, Kyutoku Y, Soya H. 2010. Ac
- ute moderate exercise elicits increased dorsolateral prefrontal activation and improves co
- gnitive performance with Stroop test. Neuroimage 50(4):1702-10. Epub 20091216. doi:
- 10.1016/j.neuroimage.2009.12.023.
- 4. Byun K, Hyodo K, Suwabe K, Ochi G, Sakairi Y, Kato M, Dan I, Soya H. 2014. Posit
- ive effect of acute mild exercise on executive function via arousal-related prefrontal acti vations: an fNIRS study. Neuroimage 98:336-45. Epub 20140502. doi: 10.1016/j.neuroi mage.2014.04.067.
- 5. Hwang J, Brothers RM, Castelli DM, Glowacki EM, Chen YT, Salinas MM, Kim J, J
- ung Y, Calvert HG. 2016. Acute high-intensity exercise-induced cognitive enhancement
- and brain-derived neurotrophic factor in young, healthy adults. Neurosci Lett 630:247-5
- 3. Epub 20160720. doi: 10.1016/j.neulet.2016.07.033.
- 6. Chang H, Kim K, Jung YJ, Kato M. 2017. Effects of acute high-Intensity resistance e xercise on cognitive function and oxygenation in prefrontal cortex. J Exerc Nutrition Bio chem 21(2):1-8. doi: 10.20463/jenb.2017.0012.
- 7. McMorris T, Sproule J, Turner A, Hale BJ. 2011. Acute, intermediate intensity exercis
- e, and speed and accuracy in working memory tasks: a meta-analytical comparison of ef
- fects. Physiol Behav 102(3-4):421-8. Epub 20101214. doi: 10.1016/j.physbeh.2010.12.0 07.
-
- 8. Valkenborghs SR, Noetel M, Hillman CH, Nilsson M, Smith JJ, Ortega FB, Lubans D
- R. 2019. The impact of physical activity on brain structure and function in youth: a syste
- matic review. Pediatrics 144(4):e20184032. doi: 10.1542/peds.2018-4032.

- 10. Miyake A, Friedman NP, Emerson MJ, Witzki AH, Howerter A, Wager TD. 2000. Th
- e unity and diversity of executive functions and their contributions to complex "frontal l
- obe" tasks: a latent variable analysis. Cogn Psychol 41(1):49-100. doi: 10.1006/cogp.19 99.0734.
- 11. Committee on Physical Activity, Physical Education in the School Environment, Foo
- d and Nutrition Board, Institute of Medicine. In: Kohl HW, III, Cook HD, editors. Educa
- ting the Student Body: Taking Physical Activity and Physical Education to School. Wash

ington (DC): National Academies Press (US); 2013. p. 259-309.

- 12. Ruhland S, Lange KW. 2021. Effect of classroom-based physical activity interventio ns on attention and on-task behavior in schoolchildren: a systematic review. Sports Med Health Sci 3(3):125-33. doi: 10.1016/j.smhs.2021.08.003.
- 13. Kibbe DL, Hackett J, Hurley M, McFarland A, Schubert KG, Schultz A, Harris S. 20
- 351 11. Ten years of TAKE 10!(®): integrating physical activity with academic concepts in e
- lementary school classrooms. Prev Med 52 Suppl 1:S43-50. Epub 20110131. doi: 10.101
- 6/j.ypmed.2011.01.025.
- 14. Torre MM, Temprado JJ. 2021. A review of combined training studies in older adults
- according to a new categorization of conventional interventions. Front Aging Neurosci
- 13:808539. Epub 20220201. doi: 10.3389/fnagi.2021.808539.
- 15. Shirai A, Wadazumi T. 2022. Effect of paprika xanthophyll supplementation on cogn
- itive improvement in a multitasking exercise: a pilot study for middle-aged and older ad
- ults. Healthcare (Basel) 10(1):81. Epub 20220101. doi: 10.3390/healthcare10010081.
- 16. Tabei K-i, Satoh M, Ogawa J-i, Tokita T, Nakaguchi N, Nakao K, Kida H, Tomimoto
- H. 2018. Cognitive function and brain atrophy predict non-pharmacological efficacy in dementia: the Mihama-Kiho scan Project2. Front Aging Neurosci 10:87. doi: 10.3389/fn agi.2018.00087.
- 17. Ali N, Tian H, Thabane L, Ma J, Wu H, Zhong Q, Gao Y, Sun C, Zhu Y, Wang T. 202
- 2. The effects of dual-task training on cognitive and physical functions in older adults wi
- th cognitive impairment; a systematic review and meta-analysis. J Prev Alzheimers Dis
- 9(2):359-70. doi: 10.14283/jpad.2022.16.
- 18. Kunzler MR, Carpes FP. 2022. Moderate intensity cycling combined with cognitive
- dual-task improves selective attention. Int J Sports Med 43(6):545-52. Epub 20220114.
- doi: 10.1055/a-1684-9151.
- 19. Zhang W, Liu H, Zhang T. 2023. Immediate and short-term effects of single-task and motor-cognitive dual-task on executive function. PLoS One 18(8):e0290171. Epub 202 30816. doi: 10.1371/journal.pone.0290171.
- 20. Wollesen B, Janssen TI, Müller H, Voelcker-Rehage C. 2022. Effects of cognitive-m otor dual task training on cognitive and physical performance in healthy children and ad olescents: a scoping review. Acta Psychol 224:103498. doi: 10.1016/j.actpsy.2022.10349 8.
- 21. Chambers ES, Bridge MW, Jones DA. 2009. Carbohydrate sensing in the human mo uth: effects on exercise performance and brain activity. J Physiol 587(Pt 8):1779-94. Epu b 20090223. doi: 10.1113/jphysiol.2008.164285.
- 22. Karayigit R, Ali A, Rezaei S, Ersoz G, Lago-Rodriguez A, Dominguez R, Naderi A.
- 2021. Effects of carbohydrate and caffeine mouth rinsing on strength, muscular enduran
- ce and cognitive performance. J Int Soc Sports Nutr 18(1):63. Epub 20210926. doi: 10.1

186/s12970-021-00462-0.

23. Hirata Y, Wadazumi T, Hamada N, Shirai A, Watanabe H. 2021. The effect of carboh

ydrate mouth rinse in isometric hand grip performance (in Japanese). J Phys Fit Sports M

- ed 70(4):269-76. doi: 10.7600/jspfsm.70.269.
- 24. Shirai A, Wadazumi T, Hirata Y, Hamada N, Hongu N. 2022. Carbohydrate mouth ri
- nse and spray improve prolonged exercise performance in recreationally trained male co
- llege students. Sports (Basel) 10(4):51. Epub 20220329. doi: 10.3390/sports10040051.
- 25. Gant N, Stinear CM, Byblow WD. 2010. Carbohydrate in the mouth immediately fa
- cilitates motor output. Brain Res 1350:151-8. doi: 10.1016/j.brainres.2010.04.004.
- 26. Imazu H, Anzai T, Denpo K, Nagayoshi M, Tagai M, Hanaie K, Takeda Y, Wadazum
- i T. 2016. "Sakai kokkara taiso" no torikumi to kouka: taiso no fukyu niyoru socialcapita
- l no jousei wo mezashite (in Japanese). The Japanese Journal for Public Health Nurse 72 (8):672-7.
- 27. Abiru M, Sakai H, Sawada Y, Yamane H. 2016. The effect of the challenging two ha
- nded rhythm tapping task to DLPFC activation. Asian J Occup Ther 12(1):75-83. doi: 10 .11596/asiajot.12.75.
- 28. Chauvigné LA, Gitau KM, Brown S. 2014. The neural basis of audiomotor entrainm
- ent: an ALE meta-analysis. Front Hum Neurosci 8:776. Epub 20140930. doi: 10.3389/fn hum.2014.00776.
- 29. Kotz SA, Ravignani A, Fitch WT. 2018. The evolution of rhythm processing. Trends Cogn Sci 22(10):896-910. doi: 10.1016/j.tics.2018.08.002.
- 30. Karvonen MJ, Kentala E, Mustala O. 1957. The effects of training on heart rate; a lo ngitudinal study. Ann Med Exp Biol Fenn 35(3):307-15.
- 31. Garber CE, Blissmer B, Deschenes MR, Franklin BA, Lamonte MJ, Lee I-M, Niema

 n DC, Swain DP, American College of Sports Medicine. 2011. Quantity and quality of e xercise for developing and maintaining cardiorespiratory, musculoskeletal, and neuromo tor fitness in apparently healthy adults: guidance for prescribing exercise. Med Sci Sport s Exerc 43(7):1334-59. doi: 10.1249/MSS.0b013e318213fefb.

 erse-Stroop effects. Behav Brain Res 290:187-96. Epub 20150505. doi: 10.1016/j.bbr.20 15.04.047.

32. Song Y, Hakoda Y. 2015. An fMRI study of the functional mechanisms of Stroop/rev

- 33. Sánchez-Cubillo I, Periáñez JA, Adrover-Roig D, Rodríguez-Sánchez JM, Ríos-Lag
- o M, Tirapu J, Barceló F. 2009. Construct validity of the Trail Making Test: role of task-
- switching, working memory, inhibition/interference control, and visuomotor abilities. J I

nt Neuropsychol Soc 15(3):438-50. doi: 10.1017/S1355617709090626.

- 34. Chang YK, Labban JD, Gapin JI, Etnier JL. 2012. The effects of acute exercise on co
- gnitive performance: a meta-analysis. Brain Res 1453:87-101. Epub 20120304. doi: 10.1
- 016/j.brainres.2012.02.068.
- 35. Rodrigues Oliveira-Silva IG, Dos Santos MPP, Learsi da Silva Santos Alves SK, Lim
- a-Silva AE, Araujo GG, Ataide-Silva T. 2022. Effect of carbohydrate mouth rinse on mu
- scle strength and muscular endurance: a systematic review with meta-analysis. Crit Rev
- Food Sci Nutr 63(27):8796-807.Epub 20220404. doi: 10.1080/10408398.2022.2057417.
- 36. McMorris T. 2021. The acute exercise-cognition interaction: from the catecholamine
- s hypothesis to an interoception model. Int J Psychophysiol 170:75-88. Epub 20211016.
- doi: 10.1016/j.ijpsycho.2021.10.005.
- 37. McMorris T. 2016. Developing the catecholamines hypothesis for the acute exercise-
- cognition interaction in humans: lessons from animal studies. Physiol Behav 165:291-9.
- doi: 10.1016/j.physbeh.2016.08.011.

38. Ogoh S. 2017. Relationship between cognitive function and regulation of cerebral bl

ood flow. J Physiol Sci 67(3):345-51.Epub 20170203. doi: 10.1007/s12576-017-0525-0.

39. Garrett J, Chak C, Bullock T, Giesbrecht B. 2024. A systematic review and Bayesian

- meta-analysis provide evidence for an effect of acute physical activity on cognition in y
- oung adults. Commun Psychol 2(1):82. doi: 10.1038/s44271-024-00124-2.
- 40. Ishihara T, Drollette ES, Ludyga S, Hillman CH, Kamijo K. 2021. The effects of acu

te aerobic exercise on executive function: a systematic review and meta-analysis of indiv

- idual participant data. Neurosci Biobehav Rev 128:258-69. Epub 20210618. doi: 10.101 6/j.neubiorev.2021.06.026.
- 41. Tombaugh TN. 2004. Trail Making Test A and B: normative data stratified by age an d education. Arch Clin Neuropsychol 19(2):203-14. doi: 10.1016/S0887-6177(03)00039 -8.
- 42. Ben Ayed I, Ammar A, Boujelbane MA, Salem A, Naija S, Amor SB, Trabelsi K, Jah
- rami H, Chtourou H, Trabelsi Y, El Massioui F. 2024. Acute effect of simultaneous exerc
- ise and cognitive tasks on cognitive functions in elderly individuals with mild cognitive
- impairment. Diseases 12(7):148. Epub 20240710. doi: 10.3390/diseases12070148.
- 43. Pellegrini-Laplagne M, Dupuy O, Sosner P, Bosquet L. 2022. Acute effect of a simul taneous exercise and cognitive task on executive functions and prefrontal cortex oxygen ation in healthy older adults. Brain Sci 12(4):455. Epub 20220328. doi: 10.3390/brainsci 12040455.
- 44. Jeukendrup AE. 2013. Oral carbohydrate rinse: placebo or beneficial? Curr Sports M ed Rep 12(4):222-7. doi: 10.1249/JSR.0b013e31829a6caa.
- 45. Pomportes L, Brisswalter J, Casini L, Hays A, Davranche K. 2017. Cognitive perfor
- mance enhancement induced by caffeine, carbohydrate and guarana mouth rinsing durin

g submaximal exercise. Nutrients 9(6):589. Epub 20170609. doi: 10.3390/nu9060589.

46. De Pauw K, Roelands B, Knaepen K, Polfliet M, Stiens J, Meeusen R. 2015. Effects

of caffeine and maltodextrin mouth rinsing on P300, brain imaging, and cognitive perfo

- rmance. J Appl Physiol (1985) 118(6):776-82. Epub 20150122. doi: 10.1152/japplphysio l.01050.2014.
- 47. Rowlatt G, Bottoms L, Edmonds CJ, Buscombe R. 2017. The effect of carbohydrate

mouth rinsing on fencing performance and cognitive function following fatigue-inducin

 g fencing. Eur J Sport Sci 17(4):433-40. Epub 20161109. doi: 10.1080/17461391.2016.1 251497.

- 48. Botvinick M, Nystrom LE, Fissell K, Carter CS, Cohen JD. 1999. Conflict monitorin
- g versus selection-for-action in anterior cingulate cortex. Nature 402(6758):179-81. doi:

10.1038/46035.

49. Turner CE, Byblow WD, Stinear CM, Gant N. 2014. Carbohydrate in the mouth enh

ances activation of brain circuitry involved in motor performance and sensory perceptio

n. Appetite 80:212-9. Epub 20140521. doi: 10.1016/j.appet.2014.05.020.

Conflicts of interest

 The authors declare no conflict of interest with any financial organization regarding the material discussed in the manuscript.

Funding

This research did not receive any specific grant from funding agencies in the public,

commercial, or not-for-profit sectors.

Authors' contributions

 All authors have made substantial contributions to the design of the study and the acquisition, analysis, and interpretation of the data. Donghyun KIM participated in

- drafting the manuscript and Tsuyoshi WADAZUMI revised it critically. All authors read
- and approved the final version of the manuscript.

Data availability statement

Data generated or analyzed during this study are provided in full within the published

article.

			Age(year) Height(cm) Weight(kg)		BMI
$Male(n=8)$	Mean	21.1	173.6	66.1	22.0
	SD	0.4	4.3	8.1	3.5
Female($n=9$)	Mean	21.1	160.4	51.9	20.1
	SD	0.3	4.1	5.4	2.0
$Total(n=17)$	Mean	21.1	166.6	58.6	21.0
	SD	0.3	7.9	9.8	2.9

Table 1. Participants' physical characteristics

Table 2. Results of analysis of variance for cognitive function tests

Note. **p<0.01, ***p<0.001; Post hoc test (Bonferroni), †: vs. CON, †<0.05

Abbreviations: TMT-B, Trail making test type B; CON, Control trial; CMR, Carbohydrate mouth rinse; DE, Dual-task exercise.

Table 3. Results of t-test for cognitive function tests

Note. *p<0.05, **p<0.01, ***p<0.001;

Abbreviations: TMT-B, Trail making test type B; CON, Control trial; CMR, Carbohydrate mouth rinse; DE, Dual-task exercise.

Figure captions

Figure 1. Motions performed during block exercise

Figure 2. Trail Making Test Type B, New Stroop test Ⅱ

A: Trail Making Test Type B (TMT-B); B: New Stroop test Ⅱ

Figure 3. Experiment protocol and cognitive task

TMT-B, Trail Making Test Type B; ST, New Stroop test Ⅱ; CMR, Carbohydrate mouth

rinse; DE, Dual-task exercise

Figure 4. Carbohydrate mouth rinse (spray method)

Figure 3

