

Abstract

Non-invasive brain stimulation improves the effects of resistance exercise in the neural system of elders with Alzheimer's disease

Silvia Teixeira de Pinho ¹, Angeliete Garcez Militão ¹, Tatiane Gomes Teixeira ²,
Maurício Rocha Calomeni ³, Daniel Almeida Marinho ⁴, Vernon Furtado da Silva ³

¹ Physical Education Department of Federal University of Rondônia, Porto Velho, Brazil

² Physical Education Department of Federal University of Rondônia, Porto Velho, Rondônia, Brazil

³ Doctor's Degree Program in Nursing and Biosciences - PPgEnfBio, Federal University of the State of Rio de Janeiro - UNIRIO, Rio de Janeiro, Brazil

⁴ Universidade da Beira Interior, CIDESD, Covilhã. Portugal

***E-mail:** mauriciocalomeni@gmail.com

Conflict of Interest: Nothing to declare. **Funding:** Coordenação de Aperfeiçoamento de Pessoal de Nível Superior (CAPES), through the Grant/Award Number: 062/2016 and process number 0011.331.0025-00.62/2014 from The Fundação Rondônia de Amparo ao Desenvolvimento das Ações Científicas e Tecnológicas e à Pesquisa do Estado de Rondônia (FAPERÓ).

Alzheimer's disease (AD) is the most common neurodegenerative disease (Raskin et al., 2015). Noninvasive brain stimulation (NIBS) is another form of cognitive therapy that has received attention recently (Xu et al., 2019; Calomeni et al., 2017). To investigate whether a combination of Physical Exercise (PMED) and non-invasive brain stimulation (NIBS) can minimize some of the neurobiological impairments caused by Alzheimer's disease (AD). Eighty-eight older adults were divided into four groups (n=22), named GAE (AD receiving PMED plus NIBS), GCP (AD receiving PMED), GCN (AD receiving NIBS), and GCS (control. No intervention). Binaural beats are used in NIBS and PMED. BDNF and Tau proteins were determined from blood samples. Check alpha band and SMR rhythms by EEG. All surveys were blinded. Nonparametric tests with 5% significance and size effect tests were performed before and after the intervention. For GCS ($p>0.05$), no differences were found for all variables, and GCN and GCS showed no improvement in inhibitory control ($p>0.05$) or working memory ($p>0.05$). GCP and GAE increased executive function ($p<0.05$), decreased dementia symptoms ($p<0.05$), exhibited brain wave modulation ($p<0.05$), increased BDNF ($p<0.0001$) and decreased TAU ($p<0.05$). <0.0001) outperforms

GAE. Physical activity benefits various aspects of cognition, such as executive function, working memory, and learning, and may stimulate structural and functional plasticity in the brain (Müller et al., 2020). Previous studies have shown that regular physical activity can reduce neurological risk factors such as Parkinson's disease (Silva et al., 2022), frailty (Silva et al., 2022), and AD (Macpherson et al., 2017) which improve the quality of life. Increased expression, secretion, and downstream signalling of neurotrophic factors (BDNF, VEGF, IGF-1) are some of the mechanisms that could explain exercise-induced neuroplasticity. This study showed that exercise-induced neuroplasticity is associated with BDNF (Murawska-Cialowicz et al., 2015). In rodents, pharmacological blocking of BDNF signalling attenuates the neuroplasticity effects of physical exercise on the brain (Vaynman et al., 2004). In addition, exercise can affect brain wave patterns associated with regional activity and directly participate in cognitive and motor processes. NIBS plus PMED improves cognitive function and minimises dementia symptoms in older adults with AD, results which may be mediated by modulating brain waves, increasing BDNF, and reducing TAU.

Keywords: Working memory, Quantitative electroencephalogram, Inhibitory control, Alzheimer's disease, Ageing, Brain stimulation, Exercise and memory.

References

- Raskin J, Cummings J, Hardy J, Schuh K, Dean R. Neurobiology of Alzheimer's Disease: Integrated Molecular, Physiological, Anatomical, Biomarker, and Cognitive Dimensions. *Curr Alzheimer Res*. Published online 2015. doi:10.2174/1567205012666150701103107
- Xu Y, Qiu Z, Zhu J, et al. The modulation effect of non-invasive brain stimulation on cognitive function in patients with mild cognitive impairment: A systematic review and meta-analysis of randomized controlled trials 11 Medical and Health Sciences 1103 Clinical Sciences 11 Medica. *BMC Neurosci*. Published online 2019. doi:10.1186/s12868-018-0484-2
- Calomeni MR, Furtado da Silva V, Velasques BB, Feijó OG, Bittencourt JM, Ribeiro de Souza e Silva AP. Modulatory Effect of Association of Brain Stimulation by Light and Binaural Beats in Specific Brain Waves. *Clin Pract Epidemiol Ment Heal*. 2017;13(1):134-144. doi:10.2174/1745017901713010134
- Silva VF da, Silva DAS, Martins PC, et al. Effect of physical exercise and noninvasive brain stimulation on cognition and dementia of elderly people with frailty : A randomized study. *Int J Imaging Syst Technol*. 2022;1(July 2021):1-12. doi:10.1002/ima.22765
- Murawska-Cialowicz E, Wojna J, Zuwała-Jagiello J. Crossfit training changes brain-derived neurotrophic factor and irisin levels at rest, after wingate and progressive tests, and improves aerobic capacity and body composition of young physically active men and women. *J Physiol Pharmacol*. Published online 2015.
- Vaynman S, Ying Z, Gomez-Pinilla F. Hippocampal BDNF mediates the efficacy of exercise on synaptic plasticity and cognition. *Eur J Neurosci*. Published online 2004. doi:10.1111/j.1460-9568.2004.03720.x

- Macpherson H, Teo WP, Schneider LA, Smith AE. A life-long approach to physical activity for brain health. *Front Aging Neurosci*. Published online 2017. doi:10.3389/fnagi.2017.00147
- Calomeni MR, Rocha JAMDS, da Silva APR, et al. Brain stimulation used as biofeedback training for recovery of motor functions deteriorated by stroke. *Arq Neuropsiquiatr*. Published online 2013. doi:S0004-282X2013000300159 [pii]
- Müller P, Duderstadt Y, Lessmann V, Müller NG. Lactate and BDNF: Key Mediators of Exercise Induced Neuroplasticity? *J Clin Med*. Published online 2020. doi:10.3390/jcm9041136