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Influence of Physical Activity on Cognitive Functions - Potential Mechanisms and Benefits

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Abstract.

Background and Study Aim. The positive impact of physical activity on the proper functioning of the musculoskeletal, respiratory and circulatory system has been demonstrated and confirmed by numerous studies. The results of many research indicate that systematic physical activity has also positive effect on functions of the central nervous system. For example, improvement of the cognitive functions level, such as memory and learning, under the influence of systematic physical training has been demonstrated. The positive effect of physical activity on the central nervous system is especially visible and widely described with regard to elderly people, who develop many adverse remodeling changes in the structure of the brain. Physical activity seems to be a useful tool that may help in limiting and reducing the effects of those changes. However, particularly interesting are the studies which show that also among young people (learning youth, university students), a positive effect of physical activity on cognitive processes is observed. Currently, several hypotheses are proposed, presenting potential mechanisms underlying the beneficial effects of physical activity on the central nervous system. The first hypothesis assumes the beneficial effect of physical activity on the expression of hippocampal genes related to synaptic plasticity. The second hypothesis assumes that physical effort *per se* is an inducer of the secretion of the growth factors (e.g., BDNF, IGF-1), which have a trophic effect on the nervous system. In addition, the results of the latest scientific studies indicate that the positive effect of physical activity on the central nervous system may be due to the action of phospholipase (Gpld-1), released to the bloodstream from the liver under the influence of physical exercise.

The aim of this work is to describe the positive impact of physical activity on the central nervous system. This work indicates that due to the influence on cognitive functions, physical activity is absolutely essential to both elderly and young people population. In addition, this paper describes potential mechanisms related to the impact of physical activity on the central nervous system.

Conclusions. The results of many modern studies have confirmed and proven the observations of the beneficial effect of physical exercise on central nervous study, including the cognitive functions. It seems necessary to

educate both young and elderly people that the proper level of physical activity is a key factor allowing to maintain both physical and mental health at an appropriate, desirable level.

Keywords: young people, physical activity, cognitive function, learning, neurogenesis

Introduction

Modern human genome has been evolutionarily programmed for an active lifestyle [1,2]. However, modern *Homo sapiens*, despite being genetically programmed and phenotypically adapted to an active lifestyle, more and more often choose a sedentary lifestyle, limiting their physical activity to the necessary minimum. Research conducted by World Health Organization (WHO) showed that more than a quarter of the world's adult population is insufficiently active. Globally, over 27,5% of adults and 81% of young people lead typically sedentary lifestyle [3,4]. Physical activity is recognized as a biological need of the body, necessary for the proper functioning of many genes [5,6]. Lack of physical effort causes changes in gene expression, and thus leads to the development of civilization diseases such as: cardiovascular diseases [7], diseases of the musculoskeletal system [8], cancer [9], diabetes [10], and consequently increases the risk of the premature death. The appropriate level of physical activity is the factor that allows to maintain both physical and mental health at a correct level [5,11,12]. Among the human population aged from 18 to 65 World Health Organization recommends at least 150–300 minutes per week of moderate-intensity aerobic physical activity or at least 75–150 minutes per week of vigorous-intensity aerobic physical activity. Additionally, WHO recommends muscle-strengthening activities at least 2 days per week [13].

Influence of physical activity on cognitive function.

It is well known and widely described that physical activity has beneficial influence on the circulatory, respiratory, musculoskeletal system and on metabolic processes. Research conducted in the last few years demonstrated that systematic physical activity has also a positive effect on the functioning of the central nervous system. For example, several studies conducted among both animal and human models indicated that systematic physical activity has positive effect on cognitive functions such as learning, memory, time of reaction or attention [14,15,16]. It is very important to indicate that the positive impact of physical activity on cognitive processes seems to be independent regardless of age. In elderly people many unfavorable changes are observed along with the advancing aging process [17,18]. These changes are usually degenerative and involve a systematic and progressive reduction in the number of nerve cells amongst brain structures [19]. Neurodegenerative processes frequently affect brain structures that play crucial role in cognitive processes, such as memory, association, learning ability, leading to their impairment [20,21]. Physical exercise seems to be a valid factor that may delay or even reduce the formation of adverse changes in the central nervous system within age. In elderly, regular exercise may be a key factor to reverse age-related declines in cognitive functions [22]. Furthermore, among older population, physical activity influences both physical and mental health and thus affects on the quality of life. Although among young healthy people there are no negative functional and structural changes in the central nervous system, it has been demonstrated that physical activity may also improve their level of cognitive abilities. The results of the studies, conducted so far, has shown that among young individuals, physical activity can intensify the process of neurogenesis, increase synaptic plasticity and stimulate gene expression for factors having a trophic effect on the nervous system [23,24,25]. Additionally, in animal model studies, it was observed that among young animals, compared to older individuals, the impact of physical activity on the formation of new nerve cells and the survival of newly formed neurons is stronger [22,26]. The role of newly formed neurons in the hippocampus is yet not fully understood. One current hypothesis proves that hippocampal neurogenesis is crucial for memory consolidation and learning [27]. This hypothesis is confirmed by the results of studies conducted on rodents, which showed that an increase in the level of hippocampal neurogenesis leads to an improvement in spatial memory [22], whereas reduced proliferation of neural stem cells is linked with age-related cognitive decline [19]. Furthermore, inhibition of neuronal precursors of cell proliferation leads to memory impairment and learning disorders [28]. Positive impact of adult neurogenesis on learning and memory processes turns out to be very important for young people, high school and university students [29,30]. WHO emphasized that physical activity enhances thinking processes,

learning, and judgment skills, ensures healthy growth and young people development as well as generally improves overall well-being [13].

Regardless of age, the beneficial impact of physical activity results from its systematic practice, which leads to an increase of neural stem cell proliferation rate [16], enhances neurite outgrowth, increases progenitor cell survival [26], and increases synaptic plasticity [31]. Additionally, it also increases the gene expression of neurotransmitters and growth factors, which are acting within the central nervous system [32].

The influence mechanisms of physical activity on the central nervous system.

The positive impact of physical activity on cognitive processes has been demonstrated both in animal and human studies. However, the mechanism through which beneficial changes in the central nervous system occur seems to be complex. So far, it has not been possible to clearly explain and describe the mechanisms responsible for the positive impact of physical activity on the central nervous system. Several hypotheses have been proposed to explain potential mechanisms underlying the beneficial effects of physical activity on postnatal neurogenesis. The first one assumes the beneficial effect of physical activity on the expression of hippocampal genes related to synaptic plasticity. This influence applies particularly to genes related with the glutamatergic and GABAergic systems [33]. Under the influence of physical effort, an increased activation of the glutamatergic system genes is observed. This system is the main excitatory neurotransmission system of the brain. Physical activity, within the stimulation of the glutamatergic system, may affect the production and functioning of new neurons within the mature brain structure. In contrast to the glutamatergic system, the GABAergic system genes is the main inhibitory system in the CNS. It has been proven that physical activity inhibits the activity of the GABAergic system [34]. The other hypothesis assumes that exercise per se is an inducer of growth factors secretion. Research indicates that physical activity affects the synthesis and the release of growth factors, which have trophic effect on the nervous system [35,36,37]. Several studies indicate that physical activity may affect on the increase of synthesis and release of vascular endothelial growth factor (VEGF), brain-derived growth factor (BDNF) and insulin-derived growth factor (IGF-1) [38,39,40]. Growth factors are the extracellular signaling molecules that promote increased cell growth and maintenance. During the development of the nervous system, they are important signals, regulating the proliferation and differentiation of stem cells and progenitor cells in the brain [41]. Among adults they play a crucial role in the processes of synaptic plasticity, learning and neurogenesis [42,43]. During mature neurogenesis, growth factors influence the proliferation, differentiation and survival of newly formed cells. The effects of growth factors are complementary, but the outcome of exercise on learning is mainly regulated by IGF-1 and BDNF, while exercise-dependent stimulation of angiogenesis and neurogenesis appears to be regulated by IGF-1 and VEGF [44]. Particularly interesting are the results of the latest studies which shown that positive effect of physical activity on the central nervous system may arise from the influence on secretion of glycosylphosphatidylinositol - specific phospholipase D (Gpld-1) released from the liver under the influence of physical exercise. In animal models, concentration of Gpld-1 increases after physical activity and correlate with the improved cognitive functions. Additionally, among elder active people concentration of Gpld-1 is higher than among the same age population leading sedentary lifestyle [45].

Conclusions

The beneficial effects that physical exercise demonstrate on the body were already noticed in early Greece, where it was believed that physical activity was an essential element of everyday life, ensuring both physical and mental health. The results of many modern studies have confirmed and proven the observations of the beneficial effect of physical exercise on the entire body, including the central nervous system. Unfortunately, the results of the majority of research show that the level of physical activity in modern societies is not sufficient. Therefore, it seems necessary to educate both young and elderly people that the proper level of physical activity is a key factor allowing to maintain both physical and mental health at an appropriate, desirable level.

Conflict of interest

The authors declared not to have a conflict of interest concerning this work, authorship, and/or publications of this paper.

References

1. Cordain L, Gotshall RW, Eaton SB, Eaton SB III. Physical activity, energy expenditure and fitness: an evolutionary perspective. *Int J Sports Med*, 1998; 19: 328-335. doi: 10.1055/s-2007-971926.
2. Eaton SB, Strassman BI, Nesse RM, Neel JV, Ewald PW, Williams GC, et al. Evolutionary health promotion. *Preventive Medicine*, 2002; 34: 109-118. DOI:10.1006/pmed.2001.0876
3. Guthold R, Stevens GA, Riley LM, Bull FC. Worldwide trends in insufficient physical activity from 2001 to 2016: a pooled analysis of 358 population-based surveys with 1.9 million participants. *Lancet Glob Health*, 2018;6(10): 1077-1086. doi: 10.1016/S2214-109X(18)30357-7.
4. Guthold R, Stevens GA, Riley LM, Bull FC. Global trends in insufficient physical activity among adolescents: a pooled analysis of 298 population-based surveys with 1.6 million participants. *Lancet Child Adolesc Health*, 2020;4(1):23-35. doi: 10.1016/S2352-4642(19)30323-2.
5. Booth FW, Vyas DR. Genes, environment, and exercise. *Adv Exp Med and Biol*, 2001; 502: 13-20. doi: 10.1007/978-1-4757-3401-0_3.
6. Booth FW, Chakravarthy MV, Spangerburg EE. Exercise and gene expression: physiological regulation of the human genome through physical activity. *J Physiol*, 2002; 543: 399-411. doi: 10.1113/jphysiol.2002.019265.
7. He J, Ogden LG, Bazzano LA, Vupputuri S, Loria C, Whelton PK. Risk factors for congestive heart failure in US men and women: NHANES I epidemiologic follow-up study. *Arch Intern Med*, 2001; 161: 996-1002. doi: 10.1001/archinte.161.7.996.
8. Kolokoltsev MN, Cieslicka M, Muszkieta R. Optimization of physical training of students of high school with regard to quantitative features muscular components of their bodies. *Physical Education of Students*. 2015; 19. doi: 10.15561/20755279.2015.0104.
9. Friedenreich CM, Bryant HE, Courneya KS. Case-control study of lifetime physical activity and breast cancer risk. *Am J Epidemiol*, 2001; 154: 336-347. doi: 10.1093/aje/154.4.336.
10. Hu FB, Sigal RJ, Rich-Edwards JW, Colditz GA, Solomon CG, Willett WC, et al. Walking compared with vigorous physical activity and the risk of type 2 diabetes in women: a prospective study. *JAMA*, 1999; 282:1433-1439. doi: 10.1001/jama.282.15.1433.
11. Callaghan P. Exercise a neglected intervention in mental health care? *J Psychiatr Ment Health Nurs*, 2004; 11: 476-483. doi: 10.1111/j.1365-2850.2004.00751.x.
12. Fox KR. The influence of physical activity on mental well-being. *Public Health Nutr*, 1999; 2: 411-418. doi: 10.1017/s1368980099000567.
13. World Health Organization. Guidelines on physical activity and sedentary behavior. Geneva. 2020. doi: 10.1136/bjsports-2020-102955.
14. Kempermann G, Kuhn HG, Gage FH. More hippocampal neurons in adult mice living in an enriched environment. *Nature*, 1997; 386: 493-495. doi: 10.1038/386493a0.
15. van Praag H, Christie BR, Sejnowski TJ, Gage FH. Running enhances neurogenesis, learning, and long-term potentiation in mice. *Proc Natl Acad Sci USA*, 1999a; 96: 13427-13431. doi: 10.1073/pnas.96.23.13427.
16. van Praag H, Kempermann G, Gage FH. Running increases cell proliferation and neurogenesis in the adult mouse dentate gyrus. *Nature Neuroscience*, 1999b; 2 : 266-270. doi: 10.1038/6368.
17. Kramer AF, Hahn S, Cohen NJ, Banich MT, Mc Auley E, Harrison CR, et al. Ageing, fitness and neurocognitive function. *Nature*, 1999; 400: 418-419. doi: 10.1038/22682.
18. Kronenberg G, Bick-Sander A, Bunk E, Wolf C, Ehninger D, Kempermann G. Physical exercise prevents age-related decline in precursor cell activity in the mouse dentate gyrus. *Neurobiol Aging*, 2006; 27: 1505-1513. doi: 10.1016/j.neurobiolaging.2005.09.016.
19. Kuhn HG, Dickinson-Anson H, Gage FH. Neurogenesis in the dentate gyrus of the adult rat: age-related decrease of neuronal progenitor proliferation. *J Neurosci*, 1996; 16: 2027-2033. doi: 10.1523/JNEUROSCI.16-06-02027.1996.
20. Salthouse T. The processing-speed theory in adult age differences in cognition. *Psychol Rev* 103: 403-428, 1996 <https://doi.org/10.1037/0033-295X.103.3.403>
21. Small SA, Tsai WY, De LaPaz R, Mayeux R, Stern Y. Imaging hippocampal function across the human life span: is memory decline normal or not? *Ann Neurol*, 2002; 51: 290-295. doi: 10.1002/ana.10105.
22. van Praag H, Shubert T, Zhao C, Gage FH. Exercise enhances learning and hippocampal neurogenesis in aged mice. *J Neurosci*, 2005; 25: 8680-8685. doi: 10.1523/JNEUROSCI.1731-05.2005.

23. Hueston CM, Cryan CM, Nolan YM. Stress and adolescent hippocampal neurogenesis: diet and exercise as cognitive modulators. *Translational Psychiatry*. 2017; 7: 1081. doi: [10.1038/tp.2017.48](https://doi.org/10.1038/tp.2017.48).
24. Hopkins ME, Nitecki R, Bucci DJ. Physical exercise during adolescence versus adulthood: differential effects on object recognition memory and brain-derived neurotrophic factor levels. *Neuroscience* 2011; 194: 84-94. doi: [10.1016/j.neuroscience.2011.07.071](https://doi.org/10.1016/j.neuroscience.2011.07.071).
25. Hillman CH, Motl RW, Pontifex MB, Posthuma D, Stubbe JH, Boomsma DI, et al. Physical Activity and Cognitive Function in a Cross-Section of Younger and Older Community-Dwelling Individuals. *Health Psychology*, 2006; 25: 678–687. doi: [10.1037/0278-6133.25.6.678](https://doi.org/10.1037/0278-6133.25.6.678).
26. Wu CW, Chang YT, Yu L, Chen H, Jen CJ, Wu SY, et al. Exercise enhances the proliferation of neural stem cells and neurite growth and survival of neuronal progenitor cells in dentate gyrus of middle-aged mice. *J Appl Physiol*, 2008; 105: 1585-1594. doi: [10.1152/jappphysiol.90775.2008](https://doi.org/10.1152/jappphysiol.90775.2008).
27. Cohen N.J, Eichenbaum, H. Memory, amnesia, and the hippocampal system. MIT Press, Cambridge.1993
28. Shors TJ, Miesegae G, Beylin A, Zhao M, Rydel T, Gould E. Neurogenesis in the adult is involved in the formation of trace memories. *Nature*, 2001; 410: 372-376, 2001 doi: [10.1038/35066584](https://doi.org/10.1038/35066584).
29. Kamijo K, Takeda Y. Regular physical activity improves executive function during task switching in young adult. *Int J Psychophysiol*. 2010; 75:304-311. doi: [10.1016/j.ijpsycho.2010.01.002](https://doi.org/10.1016/j.ijpsycho.2010.01.002).
30. Gligorska-Pluncec J, Manchevska S, Bozhinovska L. Psychomotor speed in young adults with different level of physical activity. *Med Arch*, 2010; 64: 139-143.
31. Farmer J, Zhao X, van Praag H, Wodtke K, Gage FH, Christie BR. Effects of voluntary exercise on synaptic plasticity and gene expression in the dentate gyrus of adult male Sprague-Dawley rats in vivo. *Neuroscience*, 2004; 124: 71-79. doi: [10.1016/j.neuroscience.2003.09.029](https://doi.org/10.1016/j.neuroscience.2003.09.029).
32. Cotman CW, Berchtold NC. Exercise: a behavioral intervention to enhance brain health and plasticity. *Trends Neurosci*, 2002; 25: 295-301. doi: [10.1016/s0166-2236\(02\)02143-4](https://doi.org/10.1016/s0166-2236(02)02143-4).
33. Tong L, Shen H, Perreau VM, Balazs R, Cotman CW. Effects of exercise on gene-expression profile in the rat hippocampus. *Neurobiol Dis*, 2001; 8: 1046-1056. doi: [10.1006/nbdi.2001.0427](https://doi.org/10.1006/nbdi.2001.0427).
34. Molteni R, Ying Z, Gomez-Pinilla F. Differential effects acute and chronic exercise on plasticity-related genes in the rat hippocampus revealed by microarray. *Eur J Neurosci*, 2002; 16: 1107-1116. doi: [10.1046/j.1460-9568.2002.02158.x](https://doi.org/10.1046/j.1460-9568.2002.02158.x).
35. Benoit BO, Savarese T, Joly M, Engstrom CM, Pang L, Reilly J, et al. Neurotrophin channeling of neural progenitor cell differentiation. *J Neurobiol*, 2001; 46: 265-280. [https://doi.org/10.1002/1097-4695\(200103\)46:4<265::AID-NEU1007>3.0.CO;2-B](https://doi.org/10.1002/1097-4695(200103)46:4<265::AID-NEU1007>3.0.CO;2-B)
36. Shetty AK, Turner DA. In vitro survival and differentiation of neurons derived from epidermal growth factor-responsive postnatal hippocampal stem cells: inducing effects of brain-derived neurotrophic factor. *J Neurobiol*, 1998; 35: 395-425. doi: [10.1002/\(sici\)1097-4695\(19980615\)35:4<395::aid-neu7>3.0.co;2-u](https://doi.org/10.1002/(sici)1097-4695(19980615)35:4<395::aid-neu7>3.0.co;2-u).
37. Katoh-Semba R, Asano T, Ueda H, Morishita R, Takeuchi IK, Inaguma Y, et al. Riluzole enhances expression of brain-derived neurotrophic factor with consequent proliferation of granule precursor cells in the rat hippocampus. *FASEB J*, 2002; 16: 1328-1330. doi: [10.1096/fj.02-0143fje](https://doi.org/10.1096/fj.02-0143fje). Epub 2002 Jun 21.
38. Cameron HA, Hazel TG, McKay RD. Regulation of neurogenesis by growth factors and neurotransmitters. *J Neurobiol*, 1998; 36: 287-306. DOI: [10.1002/\(sici\)1097-4695\(199808\)36:2<287::aid-neu13>3.0.co;2-b](https://doi.org/10.1002/(sici)1097-4695(199808)36:2<287::aid-neu13>3.0.co;2-b).
39. Jin K, Zhu K, Sun Y, Mao XO, Xie L, Greenberg DA. Vascular endothelial growth factor (VEGF) stimulates neurogenesis in vitro and in vivo. *Proc Natl Acad Sci*, 2002; 99: 11946-11950. doi: [10.1073/pnas.182296499](https://doi.org/10.1073/pnas.182296499).
40. Vaynman S, Ying Z, Gomez-Pinilla F. Hippocampal BDNF mediates the efficacy of exercise on synaptic plasticity and cognition. *Eur J Neurosci*, 2004; 20: 2580-2590. doi: [10.1111/j.1460-9568.2004.03720.x](https://doi.org/10.1111/j.1460-9568.2004.03720.x).
41. Calof AL. Intrinsic and extrinsic factors regulating vertebrate neurogenesis. *Curr Opin Neurobiol*, 1995; 5: 19-27. doi: [10.1016/0959-4388\(95\)80082-4](https://doi.org/10.1016/0959-4388(95)80082-4).
42. Kang H, Schuman EM. Long-lasting neurotrophin induced enhancement of synaptic transmission in the adult hippocampus. *Science*, 1995; 267: 1658-1662. doi: [10.1126/science.7886457](https://doi.org/10.1126/science.7886457).
43. Arwert LI, Deijen JB, Drent ML. The relation between insulin-like growth factor I levels and cognition in health elderly: A meta-analysis. *Growth Hormone & IGF Research*, 2005; 15: 416-422. doi: [10.1016/j.ghir.2005.09.001](https://doi.org/10.1016/j.ghir.2005.09.001).
44. Cotman CW, Berchtold NC, Christie LA. Exercise builds brain health: key roles of growth factor cascades and inflammation. *Trends Neurosci*. 2007; 30, 464–472. doi: [10.1016/j.tins.2007.06.011](https://doi.org/10.1016/j.tins.2007.06.011).
45. Horowitz AM, Bieri G, Smith LK, Sanches-Diaz CI, Schroer AB, Gontier G, et al. Blood factors transfer beneficial effects of exercise on neurogenesis and cognition to the aged brain. *Science*. 2020; 10: 167–173. doi: [10.1126/science.aaw2622](https://doi.org/10.1126/science.aaw2622).