

Promoting Cognitive Resilience through Exercise in Aging: Understanding the Mechanisms and Crafting Tailored Exercise Programs for Older Adults - A Critical Review

Ghulshan Ara¹, Syed Meeran Hasnain², Asha¹, Muhammad Haris Raza³, Maria Parvez⁴, Misbah Alnoor⁴

Lecturer, United College of Physical Therapy¹, College of Physical Therapy, Faculty of Allied Health Sciences, Ziauddin University², Senior Lecturer, College of Physical Therapy, Faculty of Allied Health Sciences, Ziauddin University³, Lecturer, College of Physical Therapy, Faculty of Allied Health Sciences, Ziauddin University⁴

Corresponding Email: gulshanqureshi2222@gmail.com

Abstract

The benefits of regular physical activity and exercise are increasingly seen as key to enhancing the brain health of the elderly. Despite this growing awareness, specific guidelines for physical activities designed to improve cognitive functions are yet to be established, and the precise mechanisms that underlie cognitive benefits from such stimulation remain unclear. This review will explore current studies on how exercise and physical activities impact cognitive abilities in individuals without cognitive deficits. We aim to pinpoint the reasons behind these benefits and offer recommendations for exercise routines that could improve cognitive functions. Attention is given to three main biological mechanisms believed to drive the cognitive improvements associated with exercise: promoting neuroplasticity and growth factors, reducing inflammation biomarkers, and regulating the hypothalamic-pituitary-adrenal axis. The concluding section details factors that enhance cognitive function in the elderly, such as the nature of exercise, its frequency and type, intensity and session length, and total workout duration.

Keywords

Brain-Derived Neurotrophic Factor, Cognition, Exercise, Physical Activity.



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Introduction

In 2017, 962 million individuals aged 60 and up were anticipated to increase globally, reaching roughly 2.1 billion by 2050¹. With an increasing number of elderly persons suffering from chronic illnesses², memory impairment expresses itself in issues with memory, language, or cognition, as well as activities necessary for daily living³. This could be due to dementia or vascular disease. Today, 35.6 million individuals worldwide have dementia. By 2030, this figure might rise to 75.6 million, making dementia one of the most pressing public health concerns⁴. MCI (mild cognitive impairment) encompasses any amount of impairment between normal cognition and progressive dementia. According to research, 60-66% of people with MCI will develop cognitive impairment at some point in their lifetimes, making attempts to reduce or stop this progression critical for both quality of life and healthcare costs⁵. Stopping Alzheimer's disease, one of the most common types of dementia, for only one year might save around US 113 billion by 2030⁶. Physical activity has proven beneficial in boosting the well-being of older brains. However, few guidelines attempt to improve cognitive abilities through physical activity; their processes are unknown. As a result, this work examines existing studies on the effects of physical activity and exercise on cognitive performance⁷⁻⁸. In this review, the potential mechanisms contributing to the advantages of exercise were explored along with workouts that could improve cognitive function for adults over 50 who are not experiencing cognitive decline. Furthermore, the three primary biological mechanisms thought to be responsible for the improved cognitive function that exercise induces in older adults were highlighted as follows: (i) increased neuroplasticity and growth factors, (ii) suppression of pro-inflammatory biomarker production, and (iii) regulation of the hypothalamic-pituitary-adrenal (HPA) axis. Exercise can improve cognition in various ways, promoting all the parameters in older individuals.

Types of Exercise

Exercise is any planned, systematic, and repetitive physical activity aimed at improving or maintaining fitness⁹. There are several categories of exercises:

- Aerobic or endurance exercises are continuous and rhythmic physical activities that increase heart rate and breathing¹⁰.
- Resistance exercises, often called strength or weight training, use resistance to contract muscles, improving strength, anaerobic endurance, and skeletal muscle growth¹¹.
- Yoga is a well-known practice involving physical postures, breathing techniques, and meditation. It emphasizes flexibility, strength, and mental well-being¹¹.
- Tai Chi is a Chinese martial art involving slow, fluid movements and coordinated breathing to improve flexibility, balance, and stress relief¹².

Effects of Physical Activity on the Growth of Brain-Derived Neural Elements and Neuroplasticity

Physical exercise has a significant impact on brain-derived neural elements and neuroplasticity development. Studies show a brain loss in grey and white matter, particularly in the prefrontal and hippocampus cortex¹³. A two-year study using magnetic resonance imaging revealed that people without dementia can nevertheless have hippocampal atrophy, which can be shown in annual examinations. Those with Alzheimer's disease see greater volume reduction¹⁴. Further analysis demonstrates that ageing is connected with decreased blood and plasma levels and lower concentrations of brain-derived neural factor (BDNF), which is critical for the synergy between exercise, health, and fitness, as evaluated by enzyme-linked immunosorbent assay¹⁵.

This article emphasizes the importance of growth-related factors such as BDNF, IGF-1, and VEGF. These chemicals are critical in improving cognitive functions in older adults through physical activity, with BDNF and IGF-1 supporting neuron survival and growth while facilitating synaptic plasticity, which aids in the formation and fortification of synapses. VEGF is essential for ensuring activated neurons receive sufficient oxygen and nutrients to meet their metabolic requirements¹⁵. There is significant evidence that exercise increases growth factors in older persons. Dinoff et al.¹⁶ did a meta-analysis of 29 studies on aerobic exercise (AE) and resistance training (RE) with 910 participants from various age groups. Regardless of the type of exercise, the findings showed an increase in BDNF levels following physical activity. Another meta-analysis of AE and RE in healthy people found that aerobic exercise significantly increased peripheral BDNF levels. In a study of older adults with mild cognitive impairment (MCI), intensive aerobic exercise dramatically increased blood levels of BDNF and IGF-1. In contrast, acute resistance exercises produced relatively moderate increases. A 2014 study of 49 sedentary women found that after 16 weeks of aerobic, resistance, and motor exercises, including twice-daily cardiovascular and resistance training, blood BDNF levels and verbal language processing speeds improved compared to those on the waitlist^{17,18}. Furthermore, an RCT involving a moderate-intensity AE program for 90 older participants over a year discovered a strong association between age and blood BDNF levels, notably in persons over 65, who saw the most significant rise in BDNF following exercise¹⁹.

Another RCT among older adults with MCI found that women improved on executive functioning tests. However, plasma BDNF levels decreased after a six-month intense aerobic training program. In contrast, the study discovered that males had greater levels of IGF-1 in their plasma and performed significantly better in executive function tests²⁰. Another year-long randomized controlled trial incorporating walking and stretching exercises among healthy seniors reported no significant variations between groups in BDNF, IGF-1, and VEGF blood concentrations. On the other hand, individuals who walked had higher growth factors, which might be linked to more robust functional connectivity between their middle and para-hippocampal gyrus, as seen using functional magnetic resonance imaging²¹. To appropriately analyze the impact of exercise on growth factors in the elderly and how it links to a rise in these markers and cognitive capacities

in this cohort, high-quality RCTs and meta-analyses that focus on measurement concerns are required.

The Effect of Exercise on Inflammation

In the elderly, pro-inflammatory signals are activated by microglia, cytokines, and other inflammatory factors. These signals can alter blood artery permeability and endothelial cell function, contributing to neurodegeneration²². Ongoing or acute inflammation can produce reactive oxygen molecules, neurotoxic chemicals, and other harmful neurotoxins, all deleterious impacts on brain health²³. Notably, the human hippocampus and basal ganglia contain more significant levels of inflammation-associated enzymes than other brain regions, rendering them more prone to inflammation-related damage²⁴. Inflammatory indicators such as C-reactive protein, interleukin (IL)-6, IL-1 beta, and tumour necrosis factor-alpha tend to rise with age. This increase in inflammatory markers may be associated with diminished cognitive abilities²¹. Chronic illnesses such as atherosclerosis, type 2 diabetes, multiple sclerosis, and dementia frequently show high levels of inflammation, emphasizing its importance to health²⁵. According to research, increased levels of inflammation are associated with smaller hippocampi and medial temporal lobes in older adults and people with type 2 diabetes. A study of 3,298 older adults found that after adjusting for education level, age, and cardiovascular risk factors, serum IL-6 levels were inversely associated with Mini-Mental State Examination scores, implying that increased inflammation directly impacts cognitive functions²⁶. Meanwhile, a study of 331 healthy older people found that individuals with the highest levels of IL-6 and CRP were at a higher risk of cognitive impairment than their less active counterparts²⁷. Positive results from 13 Randomized Controlled Trials (RCTs) show that resistance and aerobic activities can lower inflammatory markers in inactive people of all ages²⁸. Furthermore, research of senior people who engaged in sixteen weeks of aerobic, neuromotor, and strength workouts found that the exercising group had significantly lower TNF and IL-6 levels and higher peripheral brain-derived neurotrophic factors than the non-exercisers²⁹.

Subsequent analyses have shown frequent exercisers with mild cognitive impairment (MCI) demonstrated significantly enhanced executive functioning and attention levels³⁰. Aerobic exercise (AE) could trigger the release of anti-inflammatory agents that reduce the activity of pro-inflammatory cytokines in older individuals³¹. There is evidence suggesting aerobic exercise could positively affect cognitive functions. Recent studies have also found aerobic exercise helpful in lowering inflammation markers in the elderly population—a factor long known to affect cognition directly. However, the specific effects of exercise-induced changes in inflammation on cognition among seniors remain largely unexplored. As such, forthcoming studies aim to delve into these pathways to understand better how exercise influences cognition through its impact on inflammation markers³². While the current data underlines the potential benefits of aerobic exercise in reducing inflammation markers, future investigations are poised to elucidate the intricate interplay between exercise's effects on inflammation and cognitive functions in ageing individuals, particularly those impacted by exercise-related cognitive concerns.

The Transformative Power of Exercise on the Hypothalamic-Pituitary-Adrenal Axis for Boosting Cognitive Function

Exercise also improves cognitive function by influencing the HPA axis. The Hypothalamic-Pituitary-Adrenal (HPA) axis is an essential neuroendocrine system that controls the body's stress response. It affects cognition by secreting stress chemicals, particularly cortisol³³. Individuals' ability to cope with stress decreases as they age. High-stress levels can overwhelm the HPA Axis, reducing cognitive performance. Long-term stress can impair sugar tolerance as well as neuroendocrine and autonomic functions^{34,35}. Furthermore, a review of 51 research studies, including over 2,486 participants from various life stages, revealed that stressful experiences strain working memory and the ability to manage thoughts³⁶. According to both cross-sectional and longitudinal research, people who have stress-related chronic illnesses are 2.7% more likely to be diagnosed with Alzheimer's disease³⁷. A link exists between greater cortisol levels and diminishing cognitive capacities in older people³⁸. However, comprehensive research on 52 seniors found that. In contrast, elevated cortisol levels are associated with a gradual cognitive decline in those with mild cognitive impairment (MCI); this is not the case for people with normal cognitive functions³⁹. Regular exercise can lower stress responses in people of all ages by modulating catecholamine and cortisol levels³⁷. It also activates the HPA axis and increases tissue sensitivity to glucocorticoids, which helps to reduce muscular inflammation and cytokine production, hence reducing overall inflammation⁴⁰. Mental and physical activities diminish activity in the sympathetic nervous system and the HPA axis⁴¹. A review of 25 Randomized Controlled Trials in patients with chronic illnesses found that yoga practitioners had significantly better sympathetic nervous system function and HPA axis regulation (measured by cortisol levels, heart rate, and blood pressure) than control groups. However, this review only included those over the age of 50⁴².

A meta-analysis encompassing 40 studies mainly involving elderly subjects revealed that practising Tai Chi was effective in ameliorating symptoms of anxiety and depression⁴³. Conversely, an observational study, which included 42 middle-aged and elderly individuals, demonstrated that mental health benefits were significantly superior to aerobic exercise alone⁴⁴. The positive effects of body-mind coordination exercises such as Tai Chi and yoga on improving cognitive performance in seniors have also been highlighted through meta-analysis⁴⁵⁻⁴⁶. Additionally, an experimental study engaging over 118 seniors in an eight-week yoga program found that participants exhibited improved executive function and lower cortisol responses than those who did not participate⁴⁴. Research indicates that leveraging the HPA axis could significantly enhance cognitive functions. However, the effectiveness of the HPA Axis as a facilitator for cognitive improvement in older adults still needs to be examined⁴⁷. Engaging in mind-body exercises has shown promise in boosting cognitive abilities in seniors by mitigating anxiety and reinstating balance within the parasympathetic nervous system, among additional mechanisms. Therefore, there is a pressing need for randomized controlled trials (RCTs) and a meta-analysis of the acquired data to verify the relationship between mental exercises, HPA axis imbalances, cognition in the elderly, and the efficacy of this particular treatment approach.

Recommendations for Enhancing Cognitive Function in Elderly Individuals

Moderate-intensity exercise affects brain functions and cognitive capacities, including memory, attention, and executive control. The ACSM Guidelines (2009) recommend a minimum of 150 minutes of moderate aerobic activity per week (30 minutes, five days a week) or 60 minutes of severe aerobic exercise (20 minutes, three days a week). Over time, participating in resistance training programs has resulted in gains in cognitive function, particularly executive skills⁴⁸. The American College of Sports Medicine recommends that older persons incorporate resistance workouts targeting the major muscle groups into their weekly regimens twice.

Much of the research on the cognitive benefits of exercise has primarily focused on aerobic or strength training programs. Comparatively, there needs to be more investigation into the effects of multicomponent exercise regimes. Tai Chi represents such a multicomponent exercise that may enhance the link between physical activity and cognitive function in elderly populations. Nonetheless, the current body of evidence remains sparse, necessitating further studies to dissect the exercise specifics and their effectiveness on different cognitive domains⁵⁰. Beyond its physical aspects, Tai Chi encompasses activities that engage multiple cognitive functions. Results from a study that measured cognitive functions against reported participation in Tai Chi and aerobic exercises revealed that seniors engaging in both exercise types exhibited superior memory performance compared to their counterparts who were involved in only one type of exercise or none at all⁵¹. While the bulk of research has focused on aerobic endurance (AE), growing evidence supports resistance exercises (RE) and mind-body practices, such as Tai Chi or yoga, in boosting cognitive performance in older adults. Studies have even reported enhanced outcomes from a combination of these exercises. A meta-analysis conducted by Colcombe and Kramer, which reviewed 18 randomized controlled trials among inactive older participants, concluded that integrating RE with AE yielded more significant cognitive benefits than engaging in AE alone or avoiding exercise altogether⁴⁹.

Barha and their team reported similar outcomes in their meta-analysis on healthy adults engaging in two different exercise modalities. Both modalities enhanced overall cognitive performance and episodic memory more than resistance exercise (RE) alone⁴⁵. Future studies could delve into the cognitive effects of various exercise forms on older individuals, such as eccentric and concentric movements of lower and upper limbs, exercises both on land and in water, repetitive exercising that includes aquatic exercises, resistance exercises, plyometric workouts, and exercises aimed at improving motor skills like balance, agility, and proprioceptive training. In a related study, Northey and colleagues reviewed 36 RCTs through meta-analysis. They found that moderate to vigorous exercise, irrespective of the type, significantly benefitted cognitive performance in older adults, though the effects were modest⁵¹.

Conclusion

This review assesses the body of research focused on the impact of physical activity on cognitive abilities, highlighting the fundamental biological mechanisms that may be involved. Studies have

illustrated that engaging in physical activities, particularly those that incorporate multiple modes and involve mental engagement, can improve cognitive functions in the elderly. However, the evidence has notable limitations, such as a limited number of studies exploring factors that might influence cognitive outcomes in older adults and a call for more rigorous research in this domain. Further studies are needed to understand better the specific details regarding exercise regimens, types of activities, and the most beneficial environments for supporting cognitive health in this demographic.

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Conflict of Interest

None.

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AUTHORS' CONTRIBUTION

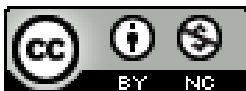
The following authors have made substantial contributions to the manuscript as under:

Conception or Design: Ara G, Hasnain SM, Asha

Acquisition, Analysis or Interpretation of Data: Ara G, Raza MH, Parvez M

Manuscript Writing & Approval: Ara G, Hasnain SM, Asha, Noor MA

All the authors agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.



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