## Exercise and cognition in older adults: is there a role for resistance training programmes?

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### ABSTRACT

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Accepted 6 November 2008 Published Online First 19 November 2008 In recent years, there has been a strong interest in physical activity as a primary behavioural prevention strategy against cognitive decline. A number of large prospective cohort studies have highlighted the protective role of regular physical activity in lowering the risk of cognitive impairment and dementia. Most prospective intervention studies of exercise and cognition to date have focused on aerobic-based exercise training. These studies highlight that aerobic-based exercise training enhances both brain structure and function. However, it has been suggested that other types of exercise training, such as resistance training, may also benefit cognition. The purpose of this brief review is to examine the evidence regarding resistance training and cognitive benefits. Three recent randomised exercise trials involving resistance training among seniors provide evidence that resistance training may have cognitive benefits. Resistance training may prevent cognitive decline among seniors via mechanisms involving insulin-like growth factor I and homocysteine. A side benefit of resistance training, albeit a very important one, is its established role in reducing morbidity among seniors. Resistance training specifically moderates the development of sarcopenia. The multifactorial deleterious sequelae of sarcopenia include increased falls and fracture risk as well as physical disability. Thus, clinicians should consider encouraging their clients to undertake both aerobic-based exercise training and resistance training not only for "physical health" but also because of the almost certain benefits for "brain health".

Cognition can be defined as "the intellectual or mental process whereby an organism becomes aware of or obtains knowledge."<sup>1</sup> Human aging is associated with declining cognition and increasing risk of dementia. Mechanisms that underlie agerelated decline in brain function include reduced volume of the cerebral white matter,<sup>2</sup> declines in the concentration, synthesis and number of receptor sites for neurotransmitters,<sup>3 4</sup> and pathological changes such as cerebral white matter lesions.<sup>5-8</sup>

The economic impact of cognitive impairment is substantial. In 2000, Alzheimer's disease and other types of dementias were the third most expensive healthcare condition in the USA, preceded by only heart disease and cancer.<sup>9</sup> Annual costs for Alzheimer's disease and other dementias were estimated at 100 billion dollars in the USA in 1997.<sup>9</sup> Strategies that would prevent the onset or progression of cognitive impairment among seniors would have enormous societal value. Brookmeyer and colleagues<sup>10</sup> estimated that, if current interventions could delay both the onset and progression of dementia by a modest 1 year, there would be nearly 9.2 million fewer cases of disease in 2050.

Physical activity has been widely promoted as a strategy for healthy aging, as it can reduce the incidence of cancer, diabetes and heart disease.<sup>11</sup> In recent years, there has been a strong interest in physical activity as a primary behavioural prevention strategy against cognitive decline. As highlighted in the review of Erickson and Kramer<sup>12</sup> in this special issue of British Journal of Sports Medicine, physical activity provides clear benefits for cognition among seniors. These neuroscientists contend that "physical activity is an inexpensive treatment that could have substantial preventative and restorative properties for cognitive and brain function."13 Certainly, a number of large prospective cohort studies have highlighted the protective role of regular physical activity in lowering the risk of cognitive impairment and dementia.<sup>14-16</sup> Most prospective intervention studies of exercise and cognition to date have focused on aerobic-based exercise training—both in animals<sup>17-20</sup> and humans.<sup>21-23</sup> These studies highlight that aerobicbased exercise training enhances both brain structure and function.<sup>17-19</sup> <sup>21-23</sup> However, it has been suggested that other types of exercise training, such as resistance training, may also benefit cognition.<sup>24</sup> The purpose of this brief review is to examine the evidence regarding resistance training and cognitive benefits.

### RESISTANCE TRAINING AND COGNITIVE OUTCOMES IN HUMANS

Although resistance training has a broad range of systemic benefits for older adults,<sup>25-29</sup> very few studies have specifically focused on its role in promoting cognitive health among seniors. Until recently, most exercise trials that examined the potential effect of resistance training on cognition in humans were limited by small sample sizes (eg, 13 and 23 participants per experimental group) and short intervention periods (eg, 8 and 16 weeks).<sup>30 31</sup>

### Possible mechanisms whereby resistance training may prevent cognitive decline

Studies with intermediate outcome measures from both human trials and laboratory experiments justify the study of resistance training and cognition. In humans, resistance training reduced serum homocysteine<sup>32</sup> and increased concentrations of insulin-like growth factor I (IGF-I).<sup>33 34</sup> Increased homocysteine concentrations are associated with impaired cognitive performance,<sup>35</sup> Alzheimer's disease<sup>36</sup> and cerebral white matter lesions.<sup>37</sup> Specifically, in a 2-year prospective study, raised

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homocysteine impaired neuropsychological functioning in otherwise cognitively intact seniors.<sup>38</sup> In rat models, raised concentrations of homocysteine are neurotoxic.<sup>39</sup> In contrast, IGF-I promotes neuronal growth, survival and differentiation and improves cognitive performance.<sup>40</sup> Thus, resistance training may prevent cognitive decline among seniors via mechanisms involving IGF-I and homocysteine.

#### Recent randomised trials of resistance training among seniors

Cassilhas and coworkers<sup>34</sup> showed that 6 months of either moderate-intensity or high-intensity resistance training significantly improved cognitive performance on standard neuropsychological tests of memory (short and long term) and verbal reasoning among senior men. They also found that serum IGF-I concentrations were higher in the resistance training groups than in the control group. Recently, Liu-Ambrose and colleagues<sup>41</sup> showed that an individualised home-based programme of balance and strength retraining, known as the Otago Exercise Program,<sup>42</sup> significantly improved executive functioning after 6 months among seniors aged 70 years and older with a recent history of falls. The finding of this study is notable given that many have hypothesised that the cognitive and neural benefits of exercise must occur within the context of social engagement for exercise to be effective.43 Details of the Otago Exercise Program are illustrated and detailed in a dedicated publication.<sup>44</sup> Briefly, basic exercises for strength and balance retraining were performed for about 20 min/day, three times a week. Adding support to these data, Brown and colleagues<sup>45</sup> showed that a 12month group-based programme of strength and balance training exercises significantly improved fluid intelligence among seniors residing in retirement villages.

Thus, there is evidence to support the hypothesis that resistance training (and balance training) may have cognitive benefits among seniors. We emphasise that resistance training may be of particular importance to senior women, as they are at greater risk of falls and fractures than senior men. However, more research is needed to clearly define the role of resistance training in the prevention of cognitive decline. Specifically, studies are needed to: (1) examine whether resistance training has similar benefits on brain function and structure to those previously shown with aerobic-based exercise training<sup>21-23</sup>; (2) examine the variables of resistance training (ie, frequency, duration and loading intensity) for maximum cognitive benefits; (3) examine whether the cognitive benefits of resistance training are limited to specific cognitive processes or general across multiple cognitive domains; (4) compare the effect of aerobic-based exercise training and resistance training on cognitive and neural plasticity. Certainly, animal studies will be essential to our understanding of the underlying mechanisms by which resistance training promotes cognitive and neural plasticity. However, for such animal studies to be conducted, methods to strength train the animals voluntarily must be developed. As highlighted by Erickson and Kramer,<sup>12</sup> there is an urgent need for research to investigate the effects of multiple types of exercise training on cognition. It is only with better understanding that we can design and deliver optimal exercise interventions to ward off cognitive decline and its associated morbidity in an aging population that is increasing yearly.

#### Additional benefits of resistance training for seniors

A very important benefit of resistance training for cognitive function is its role in reducing other morbidities among seniors. Resistance training provides a broad range of systemic benefits,<sup>25–29</sup> including moderating the development of sarcopenia—something that aerobic-based exercise training does not do. The multifactorial deleterious sequelae of sarcopenia include increased falls and fracture risk as well as physical disability.

### CLINICAL IMPLICATIONS OF CURRENT DATA ON RESISTANCE TRAINING AND COGNITION

What are the implications of these few studies of resistance training and brain function for the clinical readership of the British Journal of Sports Medicine? These findings reinforce American College of Sports Medicine (ACSM) guidelines that encourage resistance training at least twice weekly for healthy seniors.<sup>46</sup> A meta-analysis found that combined programmes of aerobic-based exercise training exercises and resistance training exercises had a greater positive effect on cognition than programmes of aerobic-based exercise training only.<sup>24</sup> Thus, clinicians should consider encouraging patients to undertake both aerobic-based exercise training and resistance training not only for "physical health" but also because of the almost certain benefits for "brain health". Many community and fitness centres have trained staff who can assist seniors to safely initiate and engage in a resistance training programme. People interested in taking up resistance training should consult their family doctor and may wish to review the ACSM position stands: exercise and physical activity in older adults<sup>47</sup> and progression models in resistance training in healthy adults.<sup>46</sup>

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#### REFERENCES

- Arwert LI, Deijen JB, Drent ML. The relation between insulin-like growth factor I levels and cognition in healthy elderly: a meta-analysis. *Growth Horm IGF Res* 2005;15:416–22.
- Haug H, Eggers R. Morphometry of the human cortex cerebri and corpus striatum during aging. *Neurobiol Aging* 1991;12:336–8; discussion 52–5.
- Adolfsson R, Gottfries CG, Roos BE, et al. Post-mortem distribution of dopamine and homovanillic acid in human brain, variations related to age, and a review of the literature. J Neural Transm 1979;45:81–105.
- de Keyser J, De Backer JP, Vauquelin G, et al. The effect of aging on the D1 dopamine receptors in human frontal cortex. Brain Res 1990;528:308–10.
- Breteler MM, van Swieten JC, Bots ML, et al. Cerebral white matter lesions, vascular risk factors, and cognitive function in a population-based study: the Rotterdam Study. Neurology 1994;44:1246–52.
- Lindgren A, Roijer A, Rudling O, et al. Cerebral lesions on magnetic resonance imaging, heart disease, and vascular risk factors in subjects without stroke. A population-based study. Stroke 1994;25:929–34.
- Garde E, Mortensen EL, Krabbe K, et al. Relation between age-related decline in intelligence and cerebral white-matter hyperintensities in healthy octogenarians: a longitudinal study. *Lancet* 2000;356:628–34.
- De Groot JC, De Leeuw FE, Oudkerk M, et al. Periventricular cerebral white matter lesions predict rate of cognitive decline. Ann Neurol 2002;52:335–41.
- Kirschstein R. Disease-specific estimates of direct and indirect costs of illness and NIH support. Fiscal year 2000 update. National Institute of Health. http://ospp.od.nih. gov/ecostudies/COIreportweb.htm (accessed 19 Nov 2008).
- Brookmeyer R, Johnson E, Ziegler-Graham K, et al. Forecasting the global burden of Alzheimer's disease. Alzheimers Dement 2007;3:186–91.
- 11. **Booth FW**, Gordon SE, Carlson CJ, *et al*. Waging war on modern chronic diseases: primary prevention through exercise biology. *J Appl Physiol* 2000;**88**:774–87.
- Erickson K, Kramer AF. Aerobic exercise effects on cognitive and neural plasticity in older adults. Br J Sports Med 2009;43:22–4.
- Kramer AF, Erickson KI. Capitalizing on cortical plasticity: influence of physical activity on cognition and brain function. *Trends Cogn Sci* 2007;11:342–8.
- Laurin D, Verreault R, Lindsay J, et al. Physical activity and risk of cognitive impairment and dementia in elderly persons. Arch Neurol 2001;58:498–504.
- Weuve J, Kang JH, Manson JE, et al. Physical activity, including walking, and cognitive function in older women. JAMA 2004;292:1454–61.
- Abbott RD, White LR, Ross GW, et al. Walking and dementia in physically capable elderly men. JAMA 2004;292:1447–53.
- van Praag H, Christie BR, Sejnowski TJ, et al. Running enhances neurogenesis, learning, and long-term potentiation in mice. Proc Natl Acad Sci USA 1999;96:13427–31.

- van Praag H, Kempermann G, Gage FH. Running increases cell proliferation and neurogenesis in the adult mouse dentate gyrus. *Nat Neurosci* 1999;2:266–70.
- Black JE, Isaacs KR, Anderson BJ, et al. Learning causes synaptogenesis, whereas motor activity causes angiogenesis, in cerebellar cortex of adult rats. Proc Natl Acad Sci USA 1990;87:5568–72.
- van Praag H. Neurogenesis and exercise: past and future directions. Neuromolecular Med 2008;10:128–40.
- Colcombe SJ, Kramer AF, Erickson KI, et al. Cardiovascular fitness, cortical plasticity, and aging. Proc Natl Acad Sci USA 2004;101:3316–21.
- Colcombe SJ, Erickson KI, Raz N, et al. Aerobic fitness reduces brain tissue loss in aging humans. J Gerontol A Biol Sci Med Sci 2003;58:176–80.
- Colcombe SJ, Erickson KI, Scalf PE, et al. Aerobic exercise training increases brain volume in aging humans. J Gerontol A Biol Sci Med Sci 2006;61:1166–70.
- Colcombe Š, Kramer AF. Fitness effects on the cognitive function of older adults: a meta-analytic study. *Psychol Sci* 2003;14:125–30.
- Borst SE. Interventions for sarcopenia and muscle weakness in older people. Age Ageing 2004;33:548–55.
- Layne JE, Nelson ME. The effects of progressive resistance training on bone density: a review. Med Sci Sports Exerc 1999;31:25–30.
- Skelton D, Young A, Greig C, et al. Effects of resistance training on strength, power, and selected functional abilities of women aged 75 and over. J Am Geriatr Soc 1995;43:1081–7.
- Taaffe D, Duret C, Wheeler S, et al. Once-weekly resistance training improves muscle strength and neuromuscular performance in older adults. J Am Geriatr Soc 1999;47:1208–14.
- Trappe S, Williamson D, Godard M. Maintenance of whole muscle strength and size following resistance training in older men. J Gerontol A Biol Sci Med Sci 2002:57:B138–43
- Dustman RE, Ruhling RO, Russell EM, et al. Aerobic exercise training and improved neuropsychological function of older individuals. Neurobiol Aging 1984;5:35–42.
- 31. **Perrig-Chiello P**, Perrig WJ, Ehrsam R, *et al*. The effects of resistance training on well-being and memory in elderly volunteers. *Age Ageing* 1998;**27**:469–75.
- 32. Vincent KR, Braith RW, Bottiglieri T, *et al*. Homocysteine and lipoprotein levels following resistance training in older adults. *Prev Cardiol* 2003;6:197–203.
- Borst SE, De Hoyos DV, Garzarella L, et al. Effects of resistance training on insulinlike growth factor-I and IGF binding proteins. Med Sci Sports Exerc 2001;33:648–53.

- Cassilhas RC, Viana VA, Grassmann V, et al. The impact of resistance exercise on the cognitive function of the elderly. Med Sci Sports Exerc 2007;39:1401–7.
- Schafer JH, Glass TA, Bolla KI, et al. Homocysteine and cognitive function in a population-based study of older adults. J Am Geriatr Soc 2005;53:381–8.
- Seshadri S, Beiser A, Selhub J, et al. Plasma homocysteine as a risk factor for dementia and Alzheimer's disease. N Engl J Med 2002;346:476–83.
- Wager TD, Rilling JK, Smith EE, et al. Placebo-induced changes in fMRI in the anticipation and experience of pain. Science 2004;303:1162–7.
- Garcia A, Haron Y, Pulman K, et al. Increases in homocysteine are related to worsening of stroop scores in healthy elderly persons: a prospective follow-up study. J Gerontol A Biol Sci Med Sci 2004;59:1323–7.
- Kruman II, Culmsee C, Chan SL, *et al*. Homocysteine elicits a DNA damage response in neurons that promotes apoptosis and hypersensitivity to excitotoxicity. *J Neurosci* 2000;20:6920–6.
- Cotman CW, Berchtold NC. Exercise: a behavioral intervention to enhance brain health and plasticity. *Trends Neurosci* 2002;25:295–301.
- Liu-Ambrose T, Donaldson MG, Ahamed Y, et al. Otago Home-Based Strength and Balance Retraining Improves Executive Functioning in Older Fallers: A Randomized Controlled Trial. J Am Geriatr Soc 2008 Sep 15 [Epub ahead of print].
- Campbell J, Robertson M, Gardner M, et al. Randomised controlled trial of a general practice programme of home based exercise to prevent falls in elderly women. BMJ 1997;315:1065–9.
- Fabel K, Kempermann G. Physical activity and the regulation of neurogenesis in the adult and aging brain. *Neuromolecular Med* 2008;10:59–66.
- Gardner MM, Buchner DM, Robertson MC, et al. Practical implementation of an exercise-based falls prevention programme. Age Ageing 2001;30:77–83.
- 45. Brown AK, Liu-Ambrose T, Tate R, et al. The effect of group-based exercise on cognitive performance and mood in seniors residing in intermediate care and self-care retirement facilities: a randomised controlled trial. Br J Sports Med 2009. In press.
- Kraemer WJ, Adams K, Cafarelli E, et al. American College of Sports Medicine position stand: progression models in resistance training for healthy adults. *Med Sci* Sports Exerc 2002;34:364–80.
- American College of Sports Medicine. Position stand: exercise and physical activity for older adults. *Med Sci Sports Exerc* 1998;30:992–1008.

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