

A Clinical Analysis of Alzheimer's Research in Neuromuscular Physical Medicine and Rehabilitation

David L Cross^{1*} and Michael B Cross²

¹Department of Applied Medicine and Rehabilitation, Indiana State University, Terre Haute, IN, USA

²Department of Orthopedic Surgery, Rush Medical Center, Chicago, IL, USA

Purpose and Hypothesis

The purpose of this commentary is to discuss the role of clinical research as an effective method for studying the comparative effects of selected therapeutic interventions, upon the rehabilitation outcome of persons with Alzheimer's disease. Furthermore it is an analysis which will endeavor to suggest that within reason, physical therapy and orthopedic practitioners can expect patients with neurological and cognitive losses, to demonstrate functional improvement [1]. Implementing reliable experimental, clinical research and quasi-experimental design methodologies will contribute to the establishment of scientifically based standard-of-care treatment protocols [2]. The construct of innovation in neuromuscular research, is an evident trend in the rehabilitation literature [3-5]. Consistent with this trend, it is hypothesized that by integrating physical therapy treatment with cognitive therapy treatment, patients with progressive neuromuscular decline will achieve better functional outcomes.

When conducting research on human subjects with the diagnosis of Alzheimer's disease, post-hoc statistical analyses of pre-to-posttest outcome measures, using established treatment modalities, would be one possible strategy for making clinical inferences from the data collected [6]. A functional research approach to the treatment of the disease is needed [7]. By utilizing basic research design methodologies and sound statistical analyses, practitioners in their daily clinical routines, can provide important answers to questions, or discover pieces of the puzzle, that might prove to be important in the treatment and cure of Alzheimer's disease [8].

Background

Decreased cognitive focus, weakness, poor appetite, changing sleep patterns, weight loss, psychosocial withdrawal, chronic fatigue, inability to plan, confusion, difficulty processing and understanding verbal conversations, and loss of memory, are just some of the early onset to mid-stage symptoms that are experienced by Alzheimer's patients [9,10]. To affect any meaningful degree of recovery for patients with the disease, it is thought that rehabilitation treatment strategies should attempt to maximize the possibility for nerve cell regeneration, oxygenation, and possible increased brainwave activity and relearning [11-14].

Based upon the literature, therapeutic exercises, both physical/aerobic and cognitive, may be important treatment interventions for neurological patients with Alzheimer's disease [15,16]. With certain forms of exercise, alternate neuromuscular pathways might be developed, and cortical nerve impulse propagation might be facilitated, thereby improving a patient's ability to translate conscious thoughts into action through "willed behavior," and thus functional rehabilitation outcomes among Alzheimer's patients, would be improved [17].

For example hydraulic biometric rehabilitation machines have been shown to be safe, simple, challenging and easy to use in the physical therapy clinic. They are especially effective for neuromuscular patients. Hydraulic exercise provides a motivational component because the resistance of the machine is directly related to the force being exerted by the patient. Also, the stationary bicycle ergometer is similarly safe

and with individualized variable resistances, which allow for patient comfort, progressive cardiovascular training and self-directed exercise programming [18].

The individualized "effort dependent" nature of hydraulic resistance is effective in facilitating coordination, proprioception, motivation, and self-directed behaviors among rehabilitation patients [19]. Cardiovascular "aerobic brain" exercise routines such as bicycle ergometers, have demonstrated both physical and psychological advantages, by developing the individual's pulmonary endurance, functional stamina, and general well-being [20]. Furthermore, "cognitive exercise" activities are important for individuals of all ages, and in particular, the multifaceted benefits derived through self-directed life-long learning include intellectual, self-esteem, and socially adaptive personality variables [21-23].

Recent experimental research findings in regenerative biology and functional kinesiology have challenged the once accepted scientific notion that neurons cannot regenerate. In fact pluripotent embryonic cells and the regeneration of nerve cells may hold hope for the future of neurological rehabilitation [24]. Physical therapy findings have suggested improved functional outcomes among neurological patients, thereby reinforcing the validity of this concept [25].

On the other hand, the nerve cell is extremely vulnerable to oxygen deprivation, and thus oxygenation and aerobic exercise appear to be essential components of neurological rehabilitation programming [26,27]. However based upon this review, ongoing Alzheimer's clinical research is warranted.

Although different physical and cognitive therapies have been employed in a variety of neurological rehabilitation settings [28], there were no articles found in the literature that studied the relative influence of adjunctive aerobic versus cognitive exercises upon the functional outcome measures of individuals with progressive cognitive loss. It would be interesting to know which such physical and/or cognitive brain exercises would be most helpful.

Future Directions

Therapeutic exercise is an important and widely used physical therapy procedure [29] and thus is routinely prescribed for patients with neuromuscular dysfunction syndromes of all types. Those with certain neurological diseases, involving both physical and cognitive

***Corresponding author:** David L Cross, Department of Applied Medicine and Rehabilitation, Indiana State University, Terre Haute, IN, USA, E-mail: David.Cross@indstate.edu

Received February 05, 2013; **Accepted** April 05, 2013; **Published** April 05, 2013

Citation: Cross DL, Cross MB (2013) A Clinical Analysis of Alzheimer's Research in Neuromuscular Physical Medicine and Rehabilitation. Int J Phys Med Rehabil 1: 116. doi:10.4172/2329-9096.1000116

Copyright: © 2013 Cross DL, et al. This is an open-access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

impairments [30], often go through different adjunctive treatments in an effort to achieve the optimal functional outcomes [31,32]. With Alzheimer's, there is an evolving body of knowledge concerned with the question "What would be the most clinically beneficial modality and medically accepted approach to treatment?"

The advantages of "physical exercise" have been well documented, including the development of muscle strength, endurance, neuromuscular proprioception, and motor control [33]. Alzheimer's patients pose unique clinical challenges. It is never known the extent to which their cognitive loss is stable, deteriorating, or if there remains a good prognosis for improvement. In addition it is not clear whether cognitive status is related to, or affected by their variable day-to-day physical function.

"Aerobic cardiovascular" and/or standard cognitive "cerebral" brain exercises, or just placebo "social interaction activities" might all be beneficial as possible adjunctive exercise treatments. It has not been determined however, which of the different "brain exercise" therapy components might be the most effective [34-36], or whether any of these adjunctive interventions would be beneficial. The goal of each and every treatment intervention would be to facilitate maximal functional independence among patients diagnosed with Alzheimer's disease.

Establishing medically accepted rehabilitation strategies for Alzheimer's patients is consistent with a fundamental principle of the new medical model, in which all individuals are given access to treatment, regardless of their diagnosis [37]. The myriad of problems associated with Alzheimer's disease are formidable. In this era of healthcare reform, and increased fiscal accountability for care, it is important that the services rendered, are clinically effective, normalizing, and promote the human dignity of the patient. It is theorized that the null would be rejected in favor of the alternative hypothesis, and that physical exercise together with certain adjunctive aerobic and cognitive brain exercise therapies would result in improved physiological and functional outcomes among Alzheimer's physical therapy patients.

Initially, when investigating the benefits or lack thereof, derived from certain interventions, a retrospective versus a priori approach would be recommended when studying the comparative treatment effects of different modalities, in patients with Alzheimer's disease [38]. To illustrate this point, the clinician might administer similar but different prescribed adjunctive exercises and the measured "within and between group differences" on selected functional outcome variables, could subsequently be analyzed. The a posteriori results and conclusions could then be inferred to the population.

Was one of the adjunctive treatment regimens more effective than the others? With Alzheimer's disease, conducting retrospective research, looking back at the data collected during the usual course of medically necessary treatment, would be one suggested approach.

Since an Institutional Review Board (IRB) has the liability of protecting the safety and human rights of subjects, they are charged with the task of determining whether subject's "informed consent," has been satisfactorily achieved. Medical ethics issues abound in studying individuals with progressive neurological cognitive impairments, and Human Subjects Review Committees cannot compromise their standards [39].

Based upon the literature, conducting innovative neuromuscular research is essential. Alzheimer's is a complex disease, often with functionally interrelated physical and cognitive symptoms, and thus to achieve maximal clinical outcomes, it is hypothesized that

physical therapy sessions should perhaps address both components simultaneously.

As practitioners, if we hope to establish scientifically-based rehabilitation options and standard-of-care treatment protocols for patients with Alzheimer's disease, we must continue to work together and collaborate inter-professionally to find a cure.

References

1. Heyn P, Abreu BC, Ottenbacher KJ (2004) The effects of exercise training on elderly persons with cognitive impairment and dementia: a meta-analysis. *Arch Phys Med Rehabil* 85: 1694-1704.
2. Campbell DT, Stanley JC (1993) *Experimental and quasi-experimental designs for research*. Rand McNally, Chicago.
3. Goodgold S, Miami S, Ule D, Johnson JR (2001) Applicability of the functional reach and timed up and go tests for elderly individuals with Alzheimer's disease: A pilot study. *J Phys and Occ Ther in Geri* 19: 21-36.
4. Tepper DE (2013) Contemporary research topics in healthcare: crowdsourcing. *PT in Motion* 5: 26-36.
5. APTA Innovation Summit (2013) Collaborative care models.
6. Epps-Streeter M, Thies WH (2012) Alzheimer's Association International Research Grant Program, *Alz and Dem J*. Chicago, Medical and Scientific.
7. Teri L, McCurry SM, Buchner DM, Logsdon RG, LaCroix AZ, et al. (1998) Exercise and activity level in Alzheimer's disease: a potential treatment focus. *J Rehabil Res Dev* 35: 411-419.
8. Di Fabio R (2012) *Essentials of rehabilitation research: A statistical guide to clinical practice*. Amer Phys Ther Assn, APTA, Washington, DC.
9. Andrade C, Radhakrishnan R (2009) The prevention and treatment of cognitive decline and dementia: An overview of recent research on experimental treatments. *Indian J Psychiatry* 51: 12-25.
10. Rolland Y, Pillard F, Klapouszczak A, Reynish E, Thomas D, et al. (2007) Exercise program for nursing home residents with Alzheimer's disease: a 1-year randomized, controlled trial. *J Am Geriatr Soc* 55: 158-165.
11. Li C, Horn JP (2005) Physiological classification of sympathetic neurons in the rat superior cervical ganglion. (Dept Neurobiology, Univ Pittsburg School of Medicine, Pittsburg). *J Neurophysiology* 9: 340-346.
12. Khachaturian ZS, Snyder PJ (2008) Roadmap for the prevention of dementia II: Leonthal symposium. *J Alz and Dem* 5: 80-85.
13. Nguyen L, Humbert S, Saudou F, Chariot A (2010) Elongator - an emerging role in neurological disorders. *Trends Mol Med* 16: 1-6.
14. Teng YD, Yu D, Ropper AE, Li J, Kabatas S, et al. (2011) Functional multipotency of stem cells: a conceptual review of neurotrophic factor-based evidence and its role in translational research. *Curr Neuropharmacol* 9: 574-585.
15. Shabani R, Gaeini AA, Nikoo MR, Nikbackt H, Sadegifar M (2010) Effect of cardiac rehabilitation program on exercise capacity in women undergoing coronary artery bypass graft in hamadan-iran. *Int J Prev Med* 1: 247-251.
16. Pitkala KH, Raivio MM, Laakkonen ML, Tilvis RS, Kautiainen H, et al. (2010) Exercise rehabilitation on home-dwelling patients with Alzheimer's disease--a randomized, controlled trial. *Study protocol. Trials* 11: 92.
17. Blankevoort CG, van Heuvelen MJ, Scherder EJ (2013) Reliability of six physical performance tests in older people with dementia. *Phys Ther* 93: 69-78.
18. Cross DL (1988) The influence of self-directed learning contracts versus passive therapy treatment-ultrasound-in a back rehabilitation program. Indiana University Doctoral Dissertation, IUPUI, Dissertation Abstracts International, Bloomington, IN, USA.
19. Indianapolis Rehabilitation Agency (2011) Physical therapy and rehabilitation counseling services, Indianapolis, IN.
20. Lavie CJ, Milani RV, Ventura HO, Messerli FH, Murgu JP (1995) Cardiac rehabilitation, exercise training, and preventive cardiology research at Ochsner Heart and Vascular Institute. *Tex Heart Inst J* 22: 44-52.
21. Eggermont LH, Gavett BE, Volkens KM, Blankevoort CG, Scherder EJ, et al. (2010) Lower-extremity function in cognitively healthy aging, mild cognitive impairment, and Alzheimer's disease. *Arch Phys Med Rehabil* 91: 584-588.

22. (2002) Assessing health status and quality-of-life instruments: attributes and review criteria. *Qual Life Res* 11: 193-205.
23. Ehlenbach WJ, Hough CL, Crane PK, Haneuse SJ, Carson SS, et al. (2010) Association between acute care and critical illness hospitalization and cognitive function in older adults. *JAMA* 303: 763-770.
24. Teng YD, Lavik EB, Qu X, Park KI, Ourednik J, et al. (2002) Functional recovery following traumatic spinal cord injury mediated by a unique polymer scaffold seeded with neural stem cells. *Proc Natl Acad Sci U S A* 99: 3024-3029.
25. Simms K, Myers C, Adams J, Hartman J, Lindsey C, et al. (2007) Exercise tolerance testing in a cardiac rehabilitation setting: an exploratory study of its safety and practicality for exercise prescription and outcome data collection. *Proc (Bayl Univ Med Cent)* 20: 344-347.
26. Gonzalez JH (2012) Autologous stem cells in newborns with oxygen deprivation. (Hospital Univ Mexico) *Clin Trials*.
27. Snyder EY, Taylor RM, Wolfe JH (1995) Neural progenitor cell engraftment corrects lysosomal storage throughout the MPS VII mouse brain. *Nature* 374: 367-370.
28. Haddadzadeh MH, Maiya AG, Padmakumar R, Shad B, Mirbolouk F (2011) Effect of exercise-based cardiac rehabilitation on ejection fraction in coronary artery disease patients: a randomized controlled trial. *Heart Views* 12: 51-57.
29. Cross DL, Cross MB (2012) Outcome based neuromuscular care: monitoring patient happiness as a possible physical therapy home program variable in Alzheimer's disease. *J Nov Physiother* 2: 1000119.
30. Brummel NE, Jackson JC, Girard TD, Pandharipande PP, Schiro E, et al. (2012) A combined early cognitive and physical rehabilitation program for people who are critically ill: the activity and cognitive therapy in the intensive care unit (ACT-ICU) trial. *Phys Ther* 92: 1580-1592.
31. Page P (2012) Current concepts in muscle stretching for exercise and rehabilitation. *Int J Sports Phys Ther* 7: 109-119.
32. Friedell M (2007) Cognitive rehabilitation for people with Alzheimer's and dementia.
33. Cross DL (1980) The influence of physical fitness training as a rehabilitation tool. *Int J Rehabil Res* 3: 163-175.
34. Kargarfard M, Rouzbehani R, Basati F (2010) Effects of exercise rehabilitation on blood pressure of patients after myocardial infarction. *Int J Prev Med* 1: 124-130.
35. Nussbaum PD (2005) *Save Your Brain*. McGraw Hill Publishing: New York & Toronto.
36. Reisberg B, Shulman MB, Torossian C, Leng L, Zhu W (2010) Outcome over seven years of healthy adults with and without subjective cognitive impairment. *Alzheimers Dement* 6: 11-24.
37. Alzheimer's Association (2013) *An Alzheimer's Disease Glossary*. National Office, Chicago, IL.
38. Cohen JM (2012) Investigator responsibilities in human subjects' research. ISU; Aug; HRP Consulting Group & IRB, Washington, DC.
39. Hammen V (2012) IRB Net Institutional review board and research methodology factors in expedited review. Indiana State University, IRB.