

Is It Safer and More Beneficial to Work Heart Failure Patients Harder? An Editorial Commentary

Address for correspondence:

Neil A. Smart, PhD
School of Science and Technology
University of New England Library
Road
Armidale,
New South Wales 2351, Australia
nsmart2@une.edu.au

Neil A. Smart, PhD; Hashbullah Ismail, MSc

School of Science and Technology (Smart), Department of Exercise Physiology (Ismail),
University of New England, Armidale, New South Wales, Australia

Introduction

Exercise training is undoubtedly beneficial to heart failure patients in terms of improved physical fitness and quality of life,^{1,2} although a clear survival benefit has yet to be demonstrated.³ The results of some previously published exercise training trials may have been affected by several factors. First, exercise adherence is often below desired targets, which may lead to much smaller improvements than expected. Second, studies may have been affected by crossover to the exercise intervention in up to one-third of sedentary controls, which was the case in the largest trial to date, HF-ACTION (Heart Failure: A Controlled Trial Investigating Outcomes of Exercise Training).³ Finally, moderate-intensity continuous exercise has historically been the cornerstone of exercise programming, although a small volume of recent work has shown high-intensity interval exercise training to be superior for eliciting improvements in peak VO_2 and systolic heart function.^{4,5}

Exercise training guidelines have emerged in the last 20 years, for both subclinical and clinical populations. The American Heart Association (AHA) recently published physical activity guidelines for people with type II diabetes,⁶ which is especially interesting as it was perhaps the first to adjust weekly exercise time according to various activity intensities. Guidelines have been developed on the relatively large volume of data from clinical exercise training trials of moderate-intensity continuous exercise (MICE).

There seem to be 3 pillars, or established reasons, why there exists an historical preference for MICE therapy in people considered to be medium to high risk for cardiovascular events. First, the stimulus from MICE is considered sufficient to stimulate health benefits. Second, the risk of serious medical events from MICE is considered acceptable, whereas intuitively high-intensity exercise is considered by many to carry a higher risk of serious illness. Third, MICE is well tolerated by most people and is not suspected to detract from exercise adherence.

Recently, there have been a number of high-intensity intermittent exercise (HIIE) studies and study protocols published in the scientific literature. A notable study of HIIE for clinical populations was Wisloff et al's work in heart failure patients,⁵ which produced unsurpassed clinical improvements. These improvements included 46% improvements in peak VO_2 , which is regarded as the best predictor of prognosis in these patients. Wisloff et al's study

was conducted in a small sample size of 3 groups of 9 patients, and therefore in light of this, many clinicians remain unconvinced of the potential benefits of HIIE or that these programs are safe and well tolerated. Wisloff et al's work was published in the number 1 ranking cardiology journal *Circulation* (citation impact 16) despite the small sample size. The underlying success of Wisloff et al's⁵ and other HIIE work is that interval exercise allows for rest periods that make it possible for patients with heart failure to perform the total work of exercise at high intensity; exercise at high intensity is the major determinant of adaptation. In Wisloff et al's work, the comparison (continuous exercise) group completed exactly the same amount of work, thus removing ambiguity over dose responses.

Interest in clinical HIIE programs is growing, and the SMARTTEX group (Controlled Study of Myocardial Recovery After Exercise Training in Heart Failure) has grant funding for a larger HIIE program for heart failure patients.⁷ In Australia, our group has recently registered an HIIE trial protocol (presented as Supporting Information in the online version of this article), which will also involve collaborative centers in Italy.⁸ Moreover, the recent AHA scientific sessions held in 2012 devoted almost an entire session to 3 HIIE studies presented as conference papers.^{9–11} The data from these abstracts and other published works suggest that HIIE is at least as effective as MICE. Despite the intuitive opinion that HIIE programs are unsafe and poorly tolerated by patients, a recent study has suggested that high-intensity interval exercise is even more enjoyable than continuous exercise at lower intensity, which has clear implications for exercise adherence.¹² Moreover, the exercise session time required to complete the same volume of energy expenditure is often less for HIIE than in MICE sessions.⁵ As such, the required weekly exercise session frequency is also likely to be lower, as the stimulus is greater with HIIE compared to MICE. The resultant shorter time commitment required from patients is likely to equate to better exercise adherence.

One recent editorial has called for high-intensity aerobic interval training to become the clinical standard in exercise therapy for heart failure patients.¹³ This may be premature but likely in the future, as clinical improvements, safety, and adherence with HIIE all seem to be comparable or superior to those observed with MICE. To our knowledge, in approximately 110 randomized, controlled trials of exercise training in heart failure (including high-intensity exercise) that have been published to date, not 1 person has died directly due to exercise training, which equates to well over 100 000

The authors have no funding, financial relationships, or conflicts of interest to disclose.

Received: June 20, 2013
Accepted: June 20, 2013

Clin. Cardiol. (in press) 1
Published online in Wiley Online Library (wileyonlinelibrary.com)
DOI:10.1002/clc.22179 © 2013 Wiley Periodicals, Inc.

patient-hours of exercise training without a death due to exercise.

The next logical step will be to examine in a large trial if HIIIE provides better exercise adherence for the heart failure population. For this reason and those stated above, we present here a study design with larger patient numbers and follow-up periods that have been trialed previously using HIIIE. Our hypothesis is to show that time-efficient HIIIE produces better adherence than MICE, which is more time demanding. We also believe we will show mortality, serious event, and hospitalization benefits as the larger sample size and extended follow-up periods are likely to provide sufficient numbers of events to detect differences between allocation groups. Such data may have important clinical implications including improving clinical outcomes and quality of life in this growing clinical population. The results from high-intensity interval training or moderate continuous training are likely to provide an objective framework for designing a more efficacious exercise training program for heart failure patients.

References

1. Piepoli, MF, Davos C, Francis DP, et al. Exercise training meta-analysis of trials in patients with chronic heart failure (ExTraMATCH). *BMJ*. 2004;328:189.
2. Smart N, Marwick TH. Exercise training for patients with heart failure: a systematic review of factors that improve mortality and morbidity. *Am J Med*. 2004;116:693–706.
3. O'Connor CM, Whellan DJ, Lee KL, et al. Efficacy and safety of exercise training in patients with chronic heart failure: HF-ACTION randomized controlled trial. *JAMA*. 2009;301:1439–1450.
4. Smart NA. Comparison of 16 weeks of continuous versus intermittent exercise training in chronic heart failure patients. *Congest Heart Fail*. 2012;18:205–211.
5. Wisloff U, Stoylen A, Loennechen JP, et al. Superior cardiovascular effect of aerobic interval training versus moderate continuous training in heart failure patients: a randomized study. *Circulation*. 2009;119:3086–3094.
6. Marwick TH, Hordern MD, Miller T, et al. Exercise training for type 2 diabetes mellitus: impact on cardiovascular risk: a scientific statement from the American Heart Association. *Circulation*. 2009;119:3244–3262.
7. Stoylen A, Conraads V, Halle M, et al. Controlled study of myocardial recovery after interval training in heart failure: SMARTEX-HF—rationale and design. *Eur J Prev Cardiol*. 2012;19:813–821.
8. Ismail H., McFarlane JR, Nojournian AH, Dieberg G, Smart NA. This work has been provisionally accepted by J Am Coll Cardiol. Heart Failure.
9. Angadi S, Mookadam F, Lee C, Bright H, Royter A, Walsh J, Zakas D, Thompson P, Gaesser G. High Intensity Interval Training Improves Diastolic Dysfunction in Patients with Heart Failure and Preserved Ejection Fraction.. *Circulation*. 2012;126:A18118.
10. Werner C, Hauser M, Meyer T, Boehm M, Laufs U. High-intensity Interval Training Reduces P53 Expression and Increases Telomerase Activity in Circulating Mononuclear Cells of Untrained Subjects. *Circulation*. 2012;126:A18220.
11. Matsuo T, Saotome K, Seino S, Shimojo N, Matsushita A, Iemitsu M, Ohshima H, Tanaka K, Mukai C. Myocardial Mass Increases with both Sprint- and Aerobic-type Interval Exercise but not with a Traditional, Continuous Aerobic Exercise in Sedentary Adults: An 8-week Randomized Intervention Study. *Circulation*. 2012;126:A10704.
12. Bartlett JD, Close GL, MacLaren DP, et al. High-intensity interval running is perceived to be more enjoyable than moderate-intensity continuous exercise: implications for exercise adherence. *J Sports Sci*. 29, 2011;6:547–553.
13. Arena R, Myers J, Forma DE, et al. Should high-intensity-aerobic interval training become the clinical standard in heart failure? *Heart Fail Rev*. 2013;18:95–105.