



Published in final edited form as:

J Appl Gerontol. 2022 February ; 41(2): 486–495. doi:10.1177/0733464820964338.

Exercise and Cognitive Training Intervention improves Self-Care, Quality of life and Functional Capacity in Persons with Heart Failure

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Abstract

This study evaluated a 12-week, home-based combined aerobic exercise (walking) and computerized cognitive training (EX/CCT) program on heart failure (HF) self-care behaviors (Self-care of HF Index [SCHFI]), disease specific quality of life (Kansas City Cardiomyopathy Questionnaire [KCCQ] and functional capacity (6-minute walk distance) compared to exercise only (EX) or a usual care attention control (AC) stretching and flexibility program. Participants (N=69) were older, predominately female (54%) and African American (55%). There was significant improvement in self-care management ($F [2,13] = 5.7, p < 0.016$), KCCQ physical limitation subscale ($F [2, 52] = 3.4, p < 0.039$) and functional capacity (336 ± 18 versus 388 ± 20 meters, $p < 0.05$) among the EX/CCT participants. The underlying mechanisms that EX and CCT targets and the optimal dose that leads to improved outcomes are needed to design effective interventions for this rapidly growing population.

Keywords

heart failure; exercise; cognitive training; self-care

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Ethical Approval

Emory University Institutional Review Board, IRB number 00060752

This study is registered at www.clinicaltrials.gov (NCT02151266)

Conflict of Interest declaration:

None of the authors have any conflict of interest with respect to the research, authorship, and/or publication of this article.

Heart failure (HF) is increasing by epidemic proportions in the United States with over 6.5 million individuals currently diagnosed, and approximately 960,000 new cases identified yearly (Benjamin, Blaha, Chiuve, Cushman, Deo et al., 2017). Among older adults, HF is the most common reason for readmission within 30 days of hospital discharge and the most costly Medicare expenditure (Heidenreich, Albert, Allen, Bluemke, Butler, Fonarow et al., 2013). Cognitive impairment (CI) is a prevalent comorbidity in HF which can have devastating effects on the ability to carry out HF self-care behaviors (Lovell, Pham, Noaman, Davis, Johnson, Ibrahim, 2019). The cognitive decline observed in HF is more pronounced than with normal aging, but lacks the symptoms of dementia, such as impaired judgment, reasoning or difficulties performing activities of daily living. Cognitive impairment is established to contribute to adverse clinical outcomes, increased rehospitalizations and higher mortality rates in persons with HF (Gelow, Mudd, Chien, Lee, 2015).

Self-care is a naturalistic decision-making process used by individuals to make behavior choices about symptom monitoring and treatment adherence that preserve physiological stability (self-care maintenance) and responding to symptoms when they occur (self-care management). These self-care behaviors are influenced by the confidence or self-efficacy an individual has to perform them (self-care confidence) (Riegel, Carlson, Moser, Sebern, Hicks & Roland, 2004; Riegel, Lee & Dickson, 2011; Riegel, Dickson & Faulkner 2016). Participation in effective self-care, along with providers following the recommended HF guidelines, are essential for successful disease management and optimal quality of life (Yancy, Jessup, Bozkurt, Butler, Casey, Colvin, et al., 2017).

Few studies have evaluated interventions to improve self-care in patients with HF and co-existing CI. Davis and colleagues (2012) evaluated a targeted self-care teaching intervention on HF self-care and knowledge in 125 patients with HF and MCI. They reported improved knowledge among the intervention participants, but self-care remained poor as indicated by Self-care in Heart Failure Index (SCHFI) subscale scores below < 70%; hospital readmission rates were unchanged (Davis, Mintzer, Dennison-Himmelfarb, Hayat, Rotman, Allen 2012).

Other studies evaluating interventions to improve cognitive function have used aerobic exercise (Gary, Paul, Corwin, Butts, Miller, Hepburn et al., 2019; Tanne, Freimark, Poreh, Merzeliak, Bruck, Hayat Schwammenthal et al., 2005; Stanek, Gunstad, Spitznagel, Waechter, Hughes, Luyster et al., 2011; Alosco, Spitznagel, Cohen, Sweet, Josephson, Hughes et al., 2014) or computerized cognitive training (Pressler, Therrien, Riley, Chou, Ronis, Koelling et al., 2011; Pressler, Titler, Koelling, Riley, Jung, Hoyland-Domenico et al., 2015). The cognitive benefits of aerobic exercise in older adults are well-established. Only recently have the benefits of exercise to improve cognitive function in HF been investigated. Three of the 4 previous exercise studies conducted in HF that included a cognitive outcome were conducted in cardiac rehabilitation settings using 36 aerobic exercise sessions over 12 to 18-weeks but none of these included CCT or self-care measures (Tanne et al., 2005; Alosco, et al., 2014; Stanek et al., 2011). Our work is the only home-based aerobic exercise and CCT program reporting cognitive outcomes in HF which showed improved verbal memory (Gary et al., 2019).

The efficacy of CCT was reviewed in a recent meta-analysis, which reported cognitive benefits in both cognitively healthy older adults and in those with mild CI (Hill, Mowszowski, Naismith, Chadwick, Valenzuela, Lampit, 2017). Among persons with HF, improved delayed recall and working memory (Pressler et al., 2011; 2015) and verbal memory (Gary et al., 2019) have been reported. While CCT has been found to be beneficial for enhancing cognitive function in older adults, its benefit in HF remains inconclusive due to the small, under powered samples in previous studies and the lack of standardized cognitive testing (Kua, Valenzuela, Dong, 2019).

Evidence suggests that the combination of exercise and CCT may act synergistically for greater cognitive benefit than either strategy alone (Barcelos, Shah, Cohen, Hogan, Mulkerin, Arciero, 2015). General cognitive function (Shatil, 2013), executive function (Heill, Schumacher, Adelsberger, Martin, Jäncke, 2013; Barcelos et al., 2015), processing speed (León, Ureña, Bolaños, Bilbao Oña, 2015) and memory performance (Zhu, Yin, Lang, He, Li, 2016) have been shown to benefit more from a combined approach. In a recent meta-analysis, (Zhu et al., 2016) the effects of CCT and physical activity interventions on cognitive function were evaluated in healthy older adults. The analysis included 20 RCTs and 2667 participants which showed a significant effect for combined cognitive and physical activity versus (active or passive) control groups. Interventions reporting the use of combined exercise and CCT are sparse and none to our knowledge except our preliminary work have been conducted in HF.

The primary aim of the study (previously published) was to evaluate the efficacy of a 12-week aerobic exercise and CCT intervention on neurocognitive outcomes compared to exercise alone or an attention control usual care group receiving a stretching and flexibility program (Gary et al., 2019). We reported improved verbal memory among participants in the combined aerobic exercise and cognitive training group, with no significant changes observed in the exercise only or attention control groups from baseline to 3-months. In the current study, we hypothesized that participants in the combined aerobic and cognitive training program would also have the greatest improvement in HF self-care behaviors, disease specific quality of life and functional capacity compared to an exercise only (EX) and stretching and flexibility attention control groups (AC) from baseline to 3-months.

Method

Study design

A three-group, pre-post randomized controlled design was used to evaluate intervention efficacy with data collected at baseline and 3-months. The protocol was approved by the Institutional Review Board and all participants provided written informed consent before participation. Recruitment and data occurred from December 2014 to October 2016 in the southeastern United States at three large heart failure clinics affiliated with an academic health science center. After baseline measurements, participants were randomized using a table of random numbers prepared by a statistician. Participants were assigned to receive a usual care attention control stretch and flexibility program (UCAC, n=19), an exercise only intervention (EX, n=29), or exercise + cognitive training (EX/CCT, n=21) for 3

months. Data collection was conducted by trained research assistants (RAs) blinded to group assignment.

Participant eligibility

Participants were screened for eligibility using an electronic medical record. To be eligible, participants had to be ambulatory, between 40-75 years of age, have a left ventricular ejection fraction (LVEF) \geq 10% documented within the last year by echocardiogram, cardiac catheterization ventriculography, or radionuclide ventriculography, stable NYHA class II-III HF, receiving medication therapy for HF according to the American College of Cardiology/American Heart Association recommendation guidelines (Yancy et al., 2017) for at least 3-months prior to study enrollment. Participants also had to be English speaking and live independently. In addition, participants had to score \geq 26 on the Montreal Cognitive Assessment (MOCA) (Nasreddine, Phillips, Bédirian, Charbonneau, Whitehead, Collin et al., 2005) and have access to a computerized device for cognitive training. Exclusions included those with unstable angina or hypertension, end-stage organ failure, and any identified or diagnosed neurological or psychological disorder that would interfere with the ability to walk or cognitive function.

Intervention

Exercise only program.—A home-based aerobic exercise program was used; participants were instructed to walk 3 times per week for 12-weeks at moderate intensity. Dose-specific exercise was based on maximum heart rate (HR) obtained during a maximal effort treadmill test. Each participant was provided a target heart rate (THR) of 60% to 70% of maximum HR achieved on the cardiopulmonary exercise test (CPET) for the exercise prescription using the heart rate reserve (heart rate maximum – resting heart rate) \times % intensity) + resting HR (Karvonen, Kentala, Mustala, 1957) calculation. A home-visit by the research team after baseline testing was conducted to provide instructions on the Polar HR monitor and pedometer (Polar HR monitor [Polar, USA; Omron pedometer) for tracking THR and steps, respectively. In addition, the participant received instructions on how to complete a written walking log to document adherence to the intervention. Under weekly telephone supervision by a member of the research team, participants began the walking sessions at 60% of THR for 30 minutes, increasing to 70% intensity and 45-minutes duration over the next 4-weeks. Participants were followed by telephone for the remainder of the 12-intervention.

Combined Exercise and Computerized Cognitive Training (EX/CCT) Program.

—The same aerobic exercise walking instruction was provided to the combined ex/cct group as discussed above in one home-based session. The Intensive Auditory Training program (formally Brain Fitness) (Posit, San Francisco, CA detailed below) computerized program was used for the cognitive training component (Posit Science). The program is based on the principles of neuroplasticity and is designed to enhance sensory integration and strengthens capability for encoding information. The Intensive Auditory program is designed to be completed over 8-weeks in 40-one-hour sessions (Pressler 2011; 2015). In a second home visit, research staff provided training and demonstrated how to use the cct program with a return demonstration provided by the participant. In the current study, all participants were

provided free access to the CCT program and completed the computerized program at home at a time convenient for them.

Usual Care Attention Control

Stretching and flexibility movements—Education, flexibility and stretching protocols were provided to controls as a time-equivalent, placebo exercise condition. Participants received a home visit to review the standardized HF educational materials. The flexibility and stretching movements were also demonstrated by the research staff during the home visits with a return demonstration by the participant. In pilot evaluations, the stretching and flexibility movements had high satisfaction and were well received but were not strong enough to influence study outcomes (Gary, Dunbar, Higgins, Corwin, Hepburn, Miller, 2018). The attention control stretching and flexibility program was delivered within 2-weeks after baseline measures were completed and participants were instructed to complete the program 2-3 times per week over the next 12-weeks. Weekly telephone calls were made for 12-weeks to discuss educational materials and to answer questions about the stretching and flexibility movements.

Adherence

To be 100% adherent for the exercise sessions, participants in the EX groups were required to document walking 3-times per week at the prescribed intensity/duration on the exercise log. Polar HR monitors and pedometers were also used to objectively document exercise intensity (i.e., maximum HR and rate of perceived exertion [RPE] achieved), duration, walking adherence and progression. Participants completed walking logs and received telephone calls weekly to discuss their walking progression.

Adherence for the Brain Fitness intervention was evaluated by the number and duration of sessions automatically logged when the participant accessed the program. Time to complete the entire program was pre-established by Posit Science to be 2310 minutes (Posit Science). Participants were considered adherent if they completed 80% or 1848 minutes of the program during the 12-week intervention period which was approximately three times per week for 180 minutes total.

Measures

Socio-demographic and clinical characteristics were validated using the electronic medical record (Table 1). The Charlson Comorbidity Index (Charlson, Pompei, Ales, MacKenzie, 1987) was used to assess the number and severity of comorbid diseases. HF etiology and type were validated in the participant's medical record.

Cognitive screening assessment—Prior to enrollment, participants were screened for global cognitive function using the Montreal Cognitive Assessment (MoCA) (Nasreddine et al., 2005). Validity of the MoCA has been previously reported in HF patients (Pressler et al., 2015). Scores range between 0 and 30, with a score of 26 to 30 considered normal or no MCI present. In the current study, participants were enrolled if they had a MoCA score of 26 and met other eligibility criteria.

Aerobic and functional capacity—A maximal symptom-limited modified balke treadmill test was used to determine aerobic capacity. The modified balke is a well-tolerated exercise test that maintains treadmill speed at 3.0 mph while advancing incline to increase workload and employed with no adverse events in our work with hf participants (Balke and Ware, 1959). Peak oxygen consumption (VO_{2peak}) was obtained using continuous gas exchange according to the American Heart Association guidelines (Gibbons, Balady, Bricker, Chaitman, Fletcher, Froelicher, et al., 1997). The 6MWT a measure of functional capacity was administered using a standardized protocol where participants walked back and forth on a 100-foot flat, level hall-way, a change of 50 meters is considered clinically significant (Rasekaba, Lee, Naughton, Williams, Holland, 2009).

HF Self-care—Self-care was measured using the SCHFI v.6.2 which is a 22-item self-report instrument that includes three sub-scales that measure the three components of HF self-care: maintenance, management and confidence (Reigel, et al., 2009; Vellone, Riegel, Cocchieri, Barbaranelli, D'Agostino, Antonetti et al., 2013). The *self-care maintenance subscale* is composed of 10 items that assesses the ability to monitor symptoms and adherence behaviors utilized to avoid worsening symptoms or a HF exacerbation (i.e., dietary salt intake and medications). The *self-care management subscale* evaluates the ability to recognize symptoms and the initiation and evaluation of treatment in response to symptoms (i.e., take an additional diuretic, consult provider, reduce fluid intake). The *self-care confidence subscale* has six items to evaluate patients' perceived ability to engage in each phase of the self-care process (e.g., preventing symptom onset and recognizing symptom changes). Raw scores are standardized from 0 to 100 with higher scores reflecting better self-care. A score of 70 on each subscale is recommended as the minimum level of self-care adequacy. The SCHFI is widely used and has undergone extensive psychometric testing and revision with excellent validity and reliability reported (Riegel, Lee, Dickson & Carlson, 2009; Vellone et al., 2013). A change in a subscale score of more than one-half of a standard deviation is considered clinically relevant.

Disease Specific Quality of Life—The Kansas City Cardiomyopathy Questionnaire (KCCQ) is a 23-item, self-report outcome measure that quantifies physical function, symptoms (frequency, severity, and recent change), social function, self-efficacy and knowledge for individuals with HF. Scores range from 0 to 100, with higher scores reflecting less disease burden, symptom severity and better quality of life. The KCCQ overall summary score includes the scales for physical limitations, symptom summary, social limitations, and quality of life (Green, Porter, Bresnahan, Spertus, 2000). The KCCQ also provides an evaluation of the magnitude of clinical change (KCCQ overall summary scores: small clinically important change, 5 points; moderate change, 10 points; and large change, 20 points or greater) (Spertus, Peterson, Conard, 2005).

Data Analysis

Baseline descriptive statistics were computed for the socio-demographic, clinical and study variables and compared across study groups. Analysis of variance (ANOVA) and chi-square tests were used to examine differences between the three study groups on all demographic, clinical and outcome variables. Linear mixed effects models were used for

testing preliminary efficacy of the intervention on the outcomes of interest over time. Separate models were fitted for individual outcomes to assess group, time and group by time interaction effects, where a significant group by time interaction is indicative of a differential effect of the intervention across groups. Analyses were first unadjusted for any baseline characteristics and then adjusted for comorbidity due to baseline differences between groups. Pairwise differences between time points were used to evaluate within group changes from baseline to 3-months and Bonferroni correction was used for multiple comparisons. Missing data patterns were assessed at each time point. An intent to treat analysis was used for all data analysis. Analyses were performed using version 24 of SPSS software.

Results

The sample comprised 69 participants who completed baseline assessments. A total of 952 individuals were screened for study participation, 807 were deemed ineligible, 76 declined and 69 were enrolled (Figure 1). Participant flow from baseline to 3-months is illustrated in Figure 1. Baseline sociodemographic and clinical characteristics are presented in Table 2. The groups were comparable at baseline with the exception of higher comorbidity in the EX only group which was controlled as a covariate in the analysis. The mean age of the sample was 61 ± 10 , the majority were female ($N=37$, 54%), African American ($N=38$, 55%) and most had some college education; mean 14 ± 3 years of education (range 8-22 years). Clinically, the majority were NYHA class II ($N= 38$, 55%), LVEF% mean was 35 ± 15 with a range of 10%–65%. LVEF, and peak $\dot{V}O_{2\text{peak}}$ ($r = -0.42$, $p = 0.002$) were significantly correlated at baseline.

A mean MoCA score of 21 ± 3 (range 13-26) indicated that many participants were experiencing mild to moderate CI at baseline with scores ≤ 16 indicating the presence of possible dementia. In the current sample, 29 (42%) participants scored ≤ 20 on the MoCA.

HF self-care

There was a significant effect of the EX/CCT intervention at 3-months on the SCHFI self-management subscale ($F [2,13] = 5.7$, $p < 0.016$); improvement also occurred in the EX only group (+ 4-points) but was not statistically significant. Self-care management scores significantly declined in the AC group from 87 to 70 ($p < 0.009$). There was a clinically meaningful increase of 5-points on the self-care maintenance subscale in the EX/CCT participants compared to no change in the EX only group and scores slightly increasing in the AC (+2 points) group. Overall, the 2 intervention groups showed greater improvement in self-care management and maintenance compared to the AC participants.

Disease Specific Quality of Life

There was a significant 13-point increase in the KCCQ physical limitation subscale ($F [2, 52] = 3.4$, $p < 0.039$) and a 5-point increase in the overall summary scores among the EX/CCT participants. There was also a loss of 12-points on the KCCQ physical limitation subscale among the AC group which suggests they perceived their physical function to worsen over 3-months and was associated with a decline in the 6MWT distance. In addition,

although not statistically significant, there was a 9-point increase on the physical limitation scale among the EX only participants. Both intervention groups had greater improvement in quality of life compared to the AC group.

The mean $\dot{V}O_{2\text{peak}}$ was 16.8 ± 5 kg/ml/min with no significant group differences occurring from bl to 3-months ($F [2, 42] = 0.13, p=0.877$). The intervention had a positive effect on 6MWT distance at 3 months ($F [2, 52] = 3.3, p=0.036$) as indicated by larger increase in the intervention groups compared to the UCAC group over time.

Exercise adherence rates

There were no significant differences in adherence rates for both exercise groups. Participants 100% adherent or walked a minimum of 3 or more days per week was 60% in the EX/CCT group and 27% were moderately adherent walking a minimum of 2 times per week. In the ex only group, the adherence rate was slightly higher with 64% walking a minimum of 3 or more days per week and 24% were moderately adherent walking a minimum of 2 days per week.

CCT adherence rates

Seventeen out of 22 participants participated in the Brain Fitness exercises. Of this number, 9 (41%) were 80% adherent completing an average of 3 days and 80 minutes total per week. Three participants were less than 80% adherent and 5 did not complete any of the Brain Fitness games.

Discussion

The current study examined the effect of a combined aerobic exercise and computerized cognitive training intervention on self-care behaviors, functional capacity and disease specific quality of life in 69 participants with stable NYHA class II and III HF over a 3-month period. At baseline, all study participants had inadequate self-care management (< 70). There was significant improvement among the combined EX/CCT group in self-care management with clinically meaningful improvements among those in the EX only group. The EX/CCT group also showed greater improvement in self-care maintenance. These findings have important clinical implications since self-care management activities such as identifying and appropriately responding to worsening HF symptoms are critically important to avoid acute exacerbations and subsequent hospitalization.

Neurogenesis and neuroplasticity are means for the brain to recover from poor perfusion and oxygen deprivation such as that occurring in hf (Vogels, et al., 2007). It is plausible that exercise and cognitive training target different mechanisms in the brain and in part explain why combination therapy may be more effective. Future research to test different exercise modalities and intensity are also warranted to identify which types of training programs are superior and may lead to better cognitive outcomes. Routine evaluation of cognitive function using simple screening tools such as the MoCA (Nasreddine et al., 2005) may assist providers to identify cognitive changes that may signal a person's difficulty in practicing effective self-care.

Patient reported outcomes are important and have been shown to predict hospitalization and death in HF (Kelkar et al., 2016). The KCCQ includes a subscale for physical limitations, which quantifies from the patient's perspective the extent to which HF symptoms influence functional capacity such as walking 1-block or climbing stairs. In the current study, EX/CCT participants perceived these functional limitations to be substantially declining at 12-weeks which was supported by a significant increase in 6-minute walk distance. A 5-point change in KCCQ overall score and 13-point increase on the physical limitation subscale was associated with a 52-meter improvement in 6-minute walk distance in the EX/CCT group which is considered clinically significant (Myers, Zaheer, Quaglietti, Madhavan, Froelicher, Heidenreich, 2006). Disease-specific measures such as the KCCQ are more sensitive to clinical changes and are particularly applicable when used in clinical practice settings (Yee, Novak, Platts, Nassif, LaRue, Vader; 2019). Most practice settings have limited time and resources so questionnaires that are brief, easy to administer and interpret are appealing. The 23-item KCCQ requires less than 10 minutes to complete and the more recent, 12-item short version, takes only 2-3 minutes without losing its validity, reliability or prognostic capabilities (Spertus & Jones, 2015). Studies support that clinicians often underestimate functional capacity among patients (Kelkar et al., 2016). Health status changes can be quantified, monitored over time and interpreted using the KCCQ. For instance, a change in score may indicate a medication adjustment is needed, identify if symptoms have worsened or if other therapies should be initiated.

The prognostic ability of the 6MW distance in HF is well established with 300 meters or less associated with higher disease burden, higher rates hospitalizations and mortality (Flynn, Lin, Moe, Howlett, Fine, Spertus et al., 2012). In the current study, 43% of the participants had a 6MW distance of 300 meters or less which is lower than previously reported (Forman, Fleg, Kitzman, Brawner, Swank, McKelvie RS et al., 2012). The reason for the lower 6MW distance may reflect demographic differences from previous studies that are predominately White males with fewer comorbidities. Our population included a higher proportion of African Americans who were female, obese who had 3 or more comorbidities which may have negatively influenced their functional capacity. The 6MWT is a simple, inexpensive and objective test that is safe in stable persons with HF in outpatient settings and may be useful for risk stratification.

An interesting finding was that all 3-study groups had adequate self-care confidence at baseline, which did not significantly change over the study duration. This is consistent with Freedland et al., (2015) who also showed no increase in self-care self-confidence among participants with HF exhibiting adequate confidence at baseline. Self-care confidence or self-efficacy was first thought to be a central component of self-care but is now thought to mediate and/or moderate self-care outcomes (Riegel & Dickson 2008). Several recent secondary analyses by Vellone and colleagues (2016) have shown that self-care confidence mediates the relationship between cognition and self-care behaviors (Vellone, Fida, D'Agostino, Mottola, Juarez-Vela, Alvaro et al., 2015). Future studies are warranted to examine longitudinal relationships between cognitive function, high and low levels of self-efficacy on performance of self-care activities and their influence on clinical outcomes.

Strengths and limitations

Findings from the current study extends research in HF to a broader population of African American females with lower global cognition and functional capacity than previously reported. Previous studies have included greater numbers of Caucasian males with higher global cognition and functional capacity (Pressler et al., 2011; 2015; Tanne 2015). Importantly, none of the participants had been previously screened for cognitive decline prior to enrolling in the study but joined the study because they perceived their memory and ability to recall information had substantially declined.

The sample size was small and not adequately powered to determine group differences; larger trials are needed to determine whether these intervention strategies are clinically beneficial. Among individuals with low global function and low literacy, cognitive reserve may have been too limited to successfully engage in the Intensive Auditory Training program. Some of our participants had limited use of computers which also may have influenced their ability and motivation to engage in this type of intervention. Although we used a placebo exercise condition that included stretching and flexibility movements, a sham computer program would have better equalized the intervention groups in relation to the CCT intervention. MOCA scores have been shown to be lower among African Americans and may not have been as reliable to measure global cognitive function in the current sample (Rossetti, Lacritz, Hyman, Cullum, Van Wright, Weiner, 2017).

In conclusion, our findings indicate that a combined EX/CCT may improve self-care management, quality of life and functional capacity in persons with HF. Because there was no significant change in the EX only group, computerized cognitive training may be a more effective intervention strategy and needs further testing with and without an exercise component in an adequately powered sample. The reasons for higher self-care management and QOL scores in the EX/CCT group are unclear but may reflect improvement in functional capacity. The prevalence of HF is anticipated to rise as the population ages, which necessitates that the management of HF include other comorbid geriatric conditions such as CI and dementia (Heidenreich, Albert, Allen, Bluemke, Butler, Fonarow et al, 2013).

Acknowledgments

Funding

Supported in part by the National Institutes of Health National Institute of Nursing Research grant no. 1P30NR014134-01 (PI-D. Waldrop-Valverde), Advancing Translational Sciences of the National Institutes of Health under award no. UL1TR000454 (D. Stephens). The content is solely the responsibility of the authors and does not necessarily represent the official views of the National Institutes of Health.

References

- Alosco ML, Spitznagel MB, Cohen R, Sweet LH, Josephson R, Hughes J, Rosneck J, Gunstad J. Cardiac rehabilitation is associated with lasting improvements in cognitive function in older adults with heart failure. *Acta Cardiol.* 2014;69(4):407–14. [PubMed: 25181916]
- Balke B & Ware RW. An experimental study of fitness of Air Force personnel. *U.S. Armed Forces Med. J* 1959;10, 675–688. [PubMed: 13659732]

- Barcelos N, Shah N, Cohen K, Hogan MJ, Mulkerrin E, Arciero PJ Aerobic and cognitive exercise (ACE) pilot study for older adults: executive function improves with cognitive challenge while exergaming. *J Int Neuropsychol Soc.* 2015; 21, 768–779. [PubMed: 26581789]
- Benjamin EJ, Blaha MJ, Chiuve SE, Cushman M, Das SR, Deo R, de Ferranti SD, Heart Disease and Stroke Statistics-2017 Update: A Report From the American Heart Association. *Circulation.* 2017;135(10):e146–e603. [PubMed: 28122885]
- Borg G Psychophysical bases of perceived exertion. *Med Sci Sports Ex.* 1982; 14:377–381.
- Charlson ME, Pompei P, Ales KL, and MacKenzie CR. A new method of classifying Prognostic comorbidity in longitudinal studies: development and validation. *J Chronic Dis.* 1987; 40(5):373–383. [PubMed: 3558716]
- Davis KK, Mintzer M, Dennison-Himmelfarb CR, Hayat MJ, Rotman S, Allen J. Targeted intervention improves knowledge but not self-care or readmissions in heart failure patients with mild cognitive impairment. *Eur J Heart Fail.* 2012;14(9):1041–9. [PubMed: 22736737]
- Flynn KE, Lin L, Moe GW, Howlett JG, Fine LJ, Spertus JA, McConnell TR, Piña IL, Weinfurt KP. Relationships between changes in patient-reported health status and functional capacity in outpatients with heart failure. *Am Heart J.* 2012;163(1):88–94. [PubMed: 22172441]
- Forman DE, Fleg JL, Kitzman DW, Brawner CA, Swank AM, McKelvie RS, Clare RM, Ellis SJ, Dunlap ME, Bittner V. 6-min walk test provides prognostic utility comparable to cardiopulmonary exercise testing in ambulatory outpatients with systolic heart failure. *J Am Coll Cardiol.* 2012; 25;60(25):2653–61. [PubMed: 23177293]
- Freedland KE, Carney RM, Rich MW, Steinmeyer BC, & Rubin EH (2015). Cognitive Behavior Therapy for Depression and Self-Care in Heart Failure Patients: A Randomized Clinical Trial. *JAMA internal medicine,* 175(11), 1773–1782. 10.1001/jamainternmed.2015.5220 [PubMed: 26414759]
- Gary RA, Paul S, Corwin E, Butts B, Miller AH, Hepburn K, Williams, Waldrop-Valverde D. Exercise and Cognitive Training as a Strategy to Improve Neurocognitive Outcomes in Heart Failure: a pilot study. *Am J Geriatric Psychiatry;* Feb 2019,
- Gary RA, Dunbar SB, Higgins MK, Corwin E, Hepburn K, Miller AH. An intervention to improve physical function and caregiving perceptions in family caregivers of persons with heart failure. *Journal of Applied Gerontology,* 2018 1:733464817746757.
- Gelow JM, Mudd JO, Chien CV, Lee CS. Usefulness of cognitive dysfunction in heart failure to predict cardiovascular risk at 180 days. *Am J Cardiol.* 2015;115(6):778–82. [PubMed: 25644853]
- Gibbons RJ, Balady GJ, Bricker JT, Chaitman BR, Fletcher GF, Froelicher VF, Mark DB, McCallister BD, Mooss AN, O'Reilly MG, Winters WL, Gibbons RJ, Antman EM, Alpert JS, Faxon DP, Fuster V, Gregoratos G, Hiratzka LF, Jacobs AK, Russell RO, Smith SC; American College of Cardiology/American Heart Association Task Force on Practice Guidelines. ACC/AHA 2002 guideline update for exercise testing: summary article. A report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (Committee to Update the 1997 Exercise Testing Guidelines). *J Am Coll Cardiol.* 2002;40(8):1531–1540. [PubMed: 12392846]
- Green CP, Porter CB, Bresnahan DR, Spertus JA. Development and evaluation of the Kansas City Cardiomyopathy Questionnaire: a new health status measure for heart failure. *J Am Coll Cardiol.* 2000;35(5):1245–55. [PubMed: 10758967]
- Heidenreich PA, Albert NM, Allen LA, Bluemke DA, Butler J, Fonarow GC, Ikonmidis JS, Khavjou O, Konstam MA, Maddox TM, Nichol G, Pham M, Piña IL, Trogon JG; American Heart Association Advocacy Coordinating Committee; Council on Arteriosclerosis, Thrombosis and Vascular Biology; Council on Cardiovascular Radiology and Intervention; Council on Clinical Cardiology; Council on Epidemiology and Prevention; Stroke Council. Forecasting the impact of heart failure in the United States: a policy statement for the American Heart Association. *Circ Heart Fail.* 2013 May;6(3):606–19. [PubMed: 23616602]
- Heill N, Schumacher V, Adelsberger R, Martin M, & Jäncke L (2013). Effects of simultaneously performed cognitive and physical training in older adults. *BMC neuroscience,* 14, 103. 10.1186/1471-2202-14-103 [PubMed: 24053148]

- Hill NT, Mowszowski L, Naismith SL, Chadwick VL, Valenzuela M, Lampit A. Computerized Cognitive Training in Older Adults With Mild Cognitive Impairment or Dementia: A Systematic Review and Meta-Analysis. *Am J Psychiatry*. 2017;174(4):329–340. [PubMed: 27838936]
- Karvonen MJ, Kentala E, Mustala O. The effects of training on heart rate: a longitudinal study. *Ann Med Exp Biol Fenn* 1957;35:307–315.
- Kelkar AA, Spertus J, Pang P, Pierson RF, Cody RJ, Pina IL, Hernandez A, Butler J. Utility of Patient-Reported Outcome Instruments in Heart Failure. *JACC Heart Fail*. 2016;4(3):165–75. doi: 10.1016/j.jchf.2015.10.015. [PubMed: 26874386]
- Kua ZJ, Valenzuela M, Dong Y. Can Computerized Cognitive Training Improve Cognition in Patients With Heart Failure?: A Review. *J Cardiovasc Nurs*. 2019;34(2):E19–E27. doi: 10.1097. [PubMed: 30585868]
- León J, Ureña A, Bolaños MJ, Bilbao A, Oña A. A combination of physical and cognitive exercise improves reaction time in persons 61–84 years old. *J Aging Phys. Act*, 2015; 23 72–77. [PubMed: 24413071]
- Lovell J, Pham T, Noaman SQ, Davis MC, Johnson M, Ibrahim JE. Self-management of heart failure in dementia and cognitive impairment: a systematic review. *BMC Cardiovasc Disord*. 2019;19(1):99. [PubMed: 31035921]
- Myers J, Zaheer N, Quaglietti S, Madhavan R, Froelicher V, Heidenreich P. Association of functional and health status measures in heart failure. *J Card Fail*. 2006;12:439–45. [PubMed: 16911910]
- Nasreddine ZS, Phillips NA, Bédirian V, Charbonneau S, Whitehead V, Collin I, Cummings JL, Chertkow H. The Montreal Cognitive Assessment, MoCA: a brief screening tool for mild cognitive impairment. *J Am Geriatrics Society*. 2005;53(4):695–699.
- Polar USA. Polar Heart Rate Monitors. <https://www.polarusa.com>
- Posit Science . Companion Guide: Using the Brain Fitness Program and Understanding the Science Behind It. Posit Science; San Francisco: 2005–2007.
- Pressler SJ, Therrien B, Riley PL, Chou CC, Ronis DL, Koelling TM, Smith DG, Sullivan BJ, Frankini AM, Giordani B. Nurse-Enhanced Memory Intervention in Heart Failure: the MEMOIR study. *J Card Fail*. 2011;17(10):832–43. [PubMed: 21962422]
- Pressler SJ, Titler M, Koelling TM, Riley PL, Jung M, Hoyland-Domenico L, Ronis DL, Smith DG, Bleske BE, Dorsey SG, Giordani B. Nurse-Enhanced Computerized Cognitive Training Increases Serum Brain-Derived Neurotrophic Factor Levels and Improves Working Memory in Heart Failure. *J Card Fail*. 2015;21(8):630–41 [PubMed: 25982826]
- Rasekaba T, Lee AL, Naughton MT, Williams TJ, Holland AE. The six-minute walk test: useful metric for the cardiopulmonary patient. *Intern Med J*. 2009;39(8):495–501 [PubMed: 19732197]
- Riegel B, Carlson B, Moser DK, Sebern M, Hicks FD, Roland V. Psychometric testing of the self-care of heart failure index. *J Card Fail*. 2004;10(4):350–60. [PubMed: 15309704]
- Riegel B, Lee CS, Dickson VV, Carlson B. An update on the self-care of heart failure index. *J Cardiovasc Nurs*. 2009;24(6):485–97. [PubMed: 19786884]
- Riegel B, Lee CS, Dickson VV. Self-care in patients with chronic heart failure. *Nat Rev Cardiol*. 2011;8(11):644–54. [PubMed: 21769111]
- Riegel B, Dickson VV. A situation-specific theory of heart failure self-care. *J Cardiovasc Nurs*. 2008;23(3):190–6. [PubMed: 18437059]
- Riegel B, Dickson VV, Faulkner KM. The Situation-Specific Theory of Heart Failure Self-Care: Revised and Updated. *J Cardiovasc Nurs*. 2016;31(3):226–35. [PubMed: 25774844]
- Riegel B, Vaughan Dickson V, Goldberg LR, Deatrck JA. Factors associated with the development of expertise in heart failure self-care. *Nurs Res*. 2007;56(4):235–43. [PubMed: 17625462]
- Rossetti HC, Lacritz LH, Hynan LS, Cullum CM, Van Wright A, Weiner MF. Montreal Cognitive Assessment Performance among Community-Dwelling African Americans. *Arch Clin Neuropsychol*. 2017;32(2):238–244. [PubMed: 28365749]
- Shatil E Does combined cognitive training and physical activity training enhance cognitive abilities more than either alone? A four-condition randomized controlled trial among healthy older adults. *Front Aging Neurosci*. 2013;5:8. [PubMed: 23531885]

- Spertus JA, Jones PG. Development and Validation of a Short Version of the Kansas City Cardiomyopathy Questionnaire. *Circ Cardiovasc Qual Outcomes*. 2015 Sep;8(5):469–76. [PubMed: 26307129]
- Spertus J, Peterson E, Conard MW, Heidenreich PA, Krumholz HM, Jones P, McCullough PA, Pina I, Tooley J, Weintraub WS, Rumsfeld JS; Cardiovascular Outcomes Research Consortium. Monitoring clinical changes in patients with heart failure: a comparison of methods. *Am Heart J*. 2005;150(4):707–15. [PubMed: 16209970]
- Stanek KM, Gunstad J, Spitznagel MB, Waechter D, Hughes JW, Luyster F, Josephson R, Rosneck J. Improvements in cognitive function following cardiac rehabilitation for older adults with cardiovascular disease. *Int J Neurosci*. 2011;121(2):86–93. [PubMed: 21062215]
- Tanne D, Freimark D, Poreh A, Merzeliak O, Bruck B, Schwammenthal Y, et al. Cognitive functions in severe congestive heart failure before and after an exercise training program. *Int J Cardiol*. (2005), 103(2), 145–149. [PubMed: 16080972]
- Theill N, Schumacher V, Adelsberger R, Martin M, Jäncke L (2013). Effects of simultaneously performed cognitive and physical training in older adults. *BMC Neurosci*. 2013;14:103. [PubMed: 24053148]
- Vellone E, Riegel B, Cocchieri A, Barbaranelli C, D'Agostino F, Antonetti G, Glaser D, Alvaro R. Psychometric testing of the Self-Care of Heart Failure Index Version 6.2. *Res Nurs Health*. 2013;36(5):500–11. [PubMed: 23832431]
- Vellone E, Fida R, D'Agostino F, Mottola A, Juarez-Vela R, Alvaro R, Riegel B. Self-care confidence may be the key: A cross-sectional study on the association between cognition and self-care behaviors in adults with heart failure. *Int J Nurs Stud*. 2015;52(11):1705–13. [PubMed: 26169451]
- Vellone E, Pancani L, Greco A, Steca P, Riegel B. Self-care confidence may be more important than cognition to influence self-care behaviors in adults with heart failure: Testing a mediation model. *Int J Nurs Stud*. 2016;60:191–9. [PubMed: 27297380]
- Vogels RL, Oosterman JM, van Harten B, Scheltens P, van der Flier WM, Schroeder-Tanka JM, Weinstein HC. Profile of cognitive impairment in chronic heart failure. *J Am Geriatr Soc*. 2007;55(11):1764–70. [PubMed: 17727641]
- Yancy Jessup, Bozkurt Butler, Casey Colvin, et al. ACC/AHA/HFSA Focused Update of the 2013 ACCF/AHA Guideline for the Management of Heart Failure: A Report of the American College of Cardiology/American Heart Association Task Force on Clinical Practice Guidelines and the Heart Failure Society of America. *Circulation*. 2017;136(6):e137–e161. [PubMed: 28455343]
- Yee D, Novak E, Platts A, Nassif ME, LaRue SJ, Vader JM. Comparison of the Kansas City Cardiomyopathy Questionnaire and Minnesota Living With Heart Failure Questionnaire in Predicting Heart Failure Outcomes. *Am J Cardiol*. 2019;123(5):807–812. [PubMed: 30587373]
- Zhu X, Yin S, Lang M, He R, Li J. The more the better? A meta-analysis on effects of combined cognitive and physical intervention on cognition in healthy older adults. *Ageing Res Rev*. 2016;31:67–79. [PubMed: 27423932]

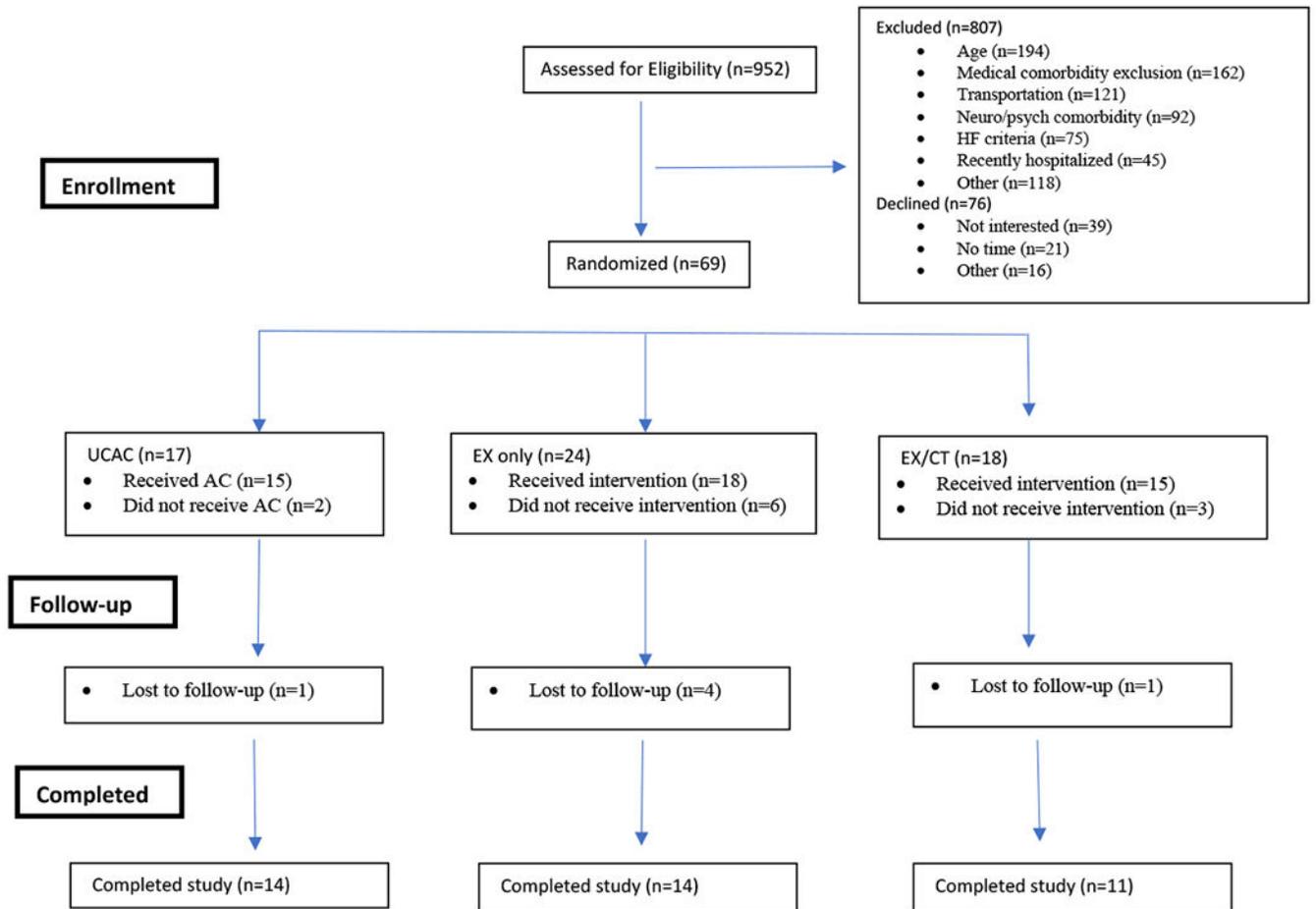


Figure 1.
CONSORT Flow Chart

Table 1.

Baseline Sociodemographic and Clinical Characteristics.

Characteristics	UCAC* (n=19)	EX only (n=29)	EX/CT (n=21)	P-value
Age, mean \pm SD	59 \pm 11	60 \pm 10	63 \pm 9	0.39
Sex, (n, %)				
Men	7 (37)	12 (41)	13 (62)	0.22
Women	12 (63)	17 (59)	8 (38)	
Race, n (%)				
African American	13 (68)	16 (55)	9 (43)	0.31
Caucasian	4 (21)	12 (41)	11 (52)	
Other	2 (11)	1 (3)	1 (5)	
Education, year, mean \pm SD	14 \pm 2	14 \pm 3	14 \pm 2	0.89
NYHA class, n (%)				
II	13 (68)	14 (48)	11 (52)	0.37
III	6 (32)	15 (42)	10 (48)	
LVEF, %, mean \pm SD	36 (14)	34 (14)	34 (19)	0.93
ICD, n (%)	7 (37)	7 (24)	11 (52)	0.81
Comorbidities, n > 3, (%)	6 (32)	12 (41)	2 (9)	0.05
BMI, mean \pm SD	31 \pm 6	32 \pm 6	32 \pm 8	0.71
Medications, n (%)				
ACEI	16 (82)	21 (73)	16 (74)	0.45
Beta-Blocker	18 (95)	28 (96)	20 (96)	0.90
ARB	11 (60)	19 (64)	14 (69)	0.49
Diuretic	16 (84)	28 (95)	20 (95)	0.09
MoCA, mean \pm SD	19 \pm 3	21 \pm 4	22 \pm 3	0.51
BDI, mean \pm SD	9 \pm 6	11 \pm 9	13 \pm 12	0.52
Peak V02 kg/mL/min, mean \pm SD	16 \pm 4	16 \pm 4	19 \pm 6	0.13
6MWT meters, mean \pm SD	336 \pm 85	359 \pm 294	319 \pm 337	0.79

* Abbreviations: UCAC= usual care attention control; EX only= exercise only; Ex/CT= exercise and cognitive training; NYHA= New York Heart Association functional class; LVEF= left ventricular ejection fraction; ICD= implantable cardiac device; BMI=Body Mass Index; ACEI= angiotensin converting enzyme inhibitor; ARB= angiotensin receptor blocker; MoCA= Montreal Cognitive Assessment; Peak V02= peak oxygen consumption; 6MWT= six minute walk test.

Table 2.

Quality of life, heart failure self-care, aerobic and functional capacity outcomes from baseline to 3-months.

Outcomes	UCAC*	Ex only	Ex/CT	RM-ANOVA Effect	3-month p-value
KCCQ (M ± SD)					
Overall summary				Time	0.509
RL	76.6 (4.5)	64.8 (3.6)	70.6 (4.3)	Time x Group	0.544
3-mos	74.6 (4.6)	67.1 (4.1)	75.0 (4.8)	Group	0.168
Physical limitations subscale					
BL	74.6 (5.8)	61.3 (4.5)	67.4 (5.4)	Time x Group	0.039
3-mos	62.6 (6.1)	70.9 (5.4)	80.5 (6.3)	Group	0.451
SCHFI (M ± SD)					
Maintenance					
BL	71.8 (4.0)	66.2 (3.2)	61.6 (3.9)	Time x Group	0.682
3-mos	74.1 (4.1)	66.7 (3.6)	66.8 (4.4)	Group	0.218
Management					
BL	87.4 (6.2)	58.7 (5.0)	58.3 (6.1)	Time x Group	0.016
3-mos	70.4 (6.2)	62.4 (5.2)	61.9 (6.2)	Group	0.035
Confidence					
BL	79.3 (4.8)	77.9 (3.8)	70.0 (4.6)	Time x Group	0.968
3-mos	78.3 (5.1)	78.1 (4.3)	70.6 (5.2)	Group	0.310
Peak V02 mL/kg/min* (M ± SD)					
BL	15.3 (1.3)	16.0 (1.0)	19.3 (1.1)	Time x Group	0.877
3-months	15.9 (1.3)	16.2 (1.0)	19.3 (1.2)	Group	0.057
6MWT (meters), (M ± SD)					
BL	324.1 (19.2)	355.1 (15.9)	337.6 (18.5)	Time x Group	0.036
3-months	311.1 (20.0)	358.8 (16.9)	388.4 (20.2)	Group	0.144

* Abbreviations: UCAC= usual care attention control; EX only= exercise only; Ex/CT= exercise and cognitive training; KCCQ= Kansas City Cardiomyopathy Questionnaire; SCHFI= Self Care Heart Failure Index; Peak V02= peak oxygen consumption; 6MWT= six minute walk test.