



Published in final edited form as:

J Psychosom Res. 2007 November ; 63(5): 463–469.

Effects of Exercise and Weight Loss on Depressive Symptoms among Men and Women with Hypertension

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Abstract

Objective—To investigate changes in depressive symptoms in hypertensive individuals participating in an exercise and weight loss intervention.

Methods—133 sedentary men and women with high blood pressure (130–180 mmHg SBP and/or 85–110 mmHg DBP) who participated in a 6 month intervention consisting of three groups: aerobic exercise, aerobic exercise and weight loss, and a waiting list control.

Results—Participants in both treatment groups demonstrated significant improvements in aerobic capacity and lower BP compared with participants in the control group. Participants in the active treatment groups who had mild to moderate depressive symptoms at baseline also exhibited greater reductions in depressive symptoms compared to participants in the control group. **Conclusion:** Results from the present study suggest that an exercise, alone or combined with weight management, may reduce self-reported depressive symptoms among patients with hypertension.

Keywords

Hypertension; depression; exercise; weight loss; aerobic fitness

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INTRODUCTION

Depression is a major health burden worldwide and is relatively common among individuals with chronic health conditions [1–3]. Although there have been important advances in pharmacological treatment of depression [4], a significant number of patients do not respond to antidepressant medication [5]. Consequently, there has been considerable interest in the development of new and effective treatments for depression. A growing body of research indicates that exercise may be a safe and effective treatment for depression [6–8] comparable to psychotherapy [9,10] and pharmacologic treatment [11,12]. The majority of previous studies have found exercise to be effective in reducing depressive symptoms both among healthy populations [7,13–17] as well as patients with heart disease [18–26]. However, it also has been noted that many studies have been plagued by methodological weaknesses, such as small samples, improper statistical analyses, and a lack of a control group, leaving uncertainty about the value of exercise as a treatment of depression [27].

The present study examines the effects of exercise on depressive symptoms among individuals with elevated blood pressure (BP). Several previous studies have shown that depression and hypertension may often exist comorbidly [28,29] and that depression may be a risk factor for the development of coronary heart disease (CHD) [30,31]. Given the increased risk of CHD among individuals with depression, interventions that reduce depressive symptoms among patients with hypertension may be especially important given the increased risk of CHD in this population. Furthermore, depressive symptoms have been associated with a greater risk for hypertension-related morbidity, cerebrovascular events, and mortality in hypertensive individuals, even at subclinical levels [32,33].

Methods

Participants

This study represents a secondary analysis of a previously reported clinical trial examining the effects of exercise and weight loss in a sample of patients with high blood pressure [34]. Briefly, participants had unmedicated high normal BP or stage 1 to 2 hypertension and were sedentary, overweight or obese, and were at least 29 years old. Exclusion criteria included current antidepressant use, current substance abuse, medical contraindications to exercise, a major psychiatric disorder requiring treatment, and ongoing participation in regular aerobic exercise. Participants were recruited from advertisements placed in local newspapers, television, and radio stations. Additionally, referrals from local clinics and BP screenings at community health fairs and local shopping centers contributed to participant recruitment.

As reported previously, 2,399 individuals were initially screened through a preliminary telephone conversation and 320 individuals underwent a subsequent medical evaluation (Figure 1). One hundred thirty three individuals met the study criteria and were randomized following a 2:2:1 schema to one of three groups for the 6-month treatment program: aerobic exercise (EX; N=54), weight management including aerobic exercise (WM; N=55), and wait list control (WLC; N=24). All participants who consented to participate in the trial were eligible for assignment to any one of the three treatment groups.

Interventions

Aerobic Exercise Only participants exercised 3 to 4 times per week at a level of 70% to 85% of their initial heart rate (HR) reserve [35] determined at the time of their baseline treadmill test. The exercise program consisted of 10 minutes of warm-up exercises, 35 minutes of cycle ergometry and walking (and eventually jogging), and 10 minutes of cool-down exercises. Participants were asked to maintain their pre-treatment diets.

Weight Management participants exercised 3 to 4 times per week using the identical protocol as previously described. In addition, participants took part in a weight management program in small groups of 3 to 4 members based on the LEARN manual [36]. The primary goal of the intervention was a weight loss of 0.5 to 1.0 kg/wk, achieved gradually by decreasing caloric consumption and fat intake through lifestyle changes.

The group format consisted of 26 weekly group sessions. Weight was recorded at each session and participants' food diaries and behavioral modification targets were reviewed. Group participation was encouraged during this process in supporting fellow group members and in problem solving around obstacles and lapses that they may have encountered. During the last 6 weeks of the program, sessions focused increasingly on weight maintenance, and group members worked on individualized plans for maintaining the changes they had made during the past 6 months.

Participants in both the EX and WM groups were instructed in how to monitor their radial pulses and maintain their heart rates (HRs) at, or above, their target level for at least 30 minutes. A trained exercise physiologist supervised all exercise sessions, and performed 2 to 3 random checks of HRs per session to ensure that participants were exercising at a sufficient intensity.

Waiting List Control participants were asked to maintain their usual dietary and exercise habits for 6 months until they were reexamined. On completion of their posttreatment assessment, participants were then allowed to select either of the 2 active treatments and could participate under supervision for 6 months.

Depression Measures

In order to investigate the effects of the intervention in alleviating depressive symptoms, participants completed the Beck Depression Inventory (BDI) [37] at baseline and after 6 months. The BDI is a standardized 21-item self-report questionnaire consisting of symptoms and attitudes related to depression including items such as self-dislike, suicidal ideation, insomnia, and sadness. The items are summed with total scores ranging from 0 to 63 with higher scores indicating higher levels depression. The BDI has been shown to be a valid and reliable measure of depression severity and a previous meta-analysis have shown the internal consistency of the BDI to yield a mean coefficient α of .86 for psychiatric patients [38].

Peak Oxygen Consumption

Maximal exercise testing was performed using the Duke-Wake Forest protocol in which graded exercise began at 3.2 kilometers per hour and 0% grade and workload was increased at a rate of 1 metabolic equivalent per minute (oxygen, 3.5 mL/kg per minute) [22]. Expired gases were collected for the determination of peak oxygen consumption using a metabolic cart (model 2900; Sensormedics, Yorba Linda, Calif). Assessments were made at baseline and again following 6 months of treatment.

Weight and Body Composition Assessment

Weight was measured by a standard balance scale. Body fat measurements were performed using a bioelectrical impedance analyzer (BIA-101Q; Quantum, Highland Heights, Ohio) in conjunction with bioelectrical impedance analyzer interpretation software (RJL Systems, Inc, Clinton Township, Mich). Measurements were done using standard right-sided, tetrapolar electrode placement with each subject in a supine position. Studies were conducted between 3 and 5pm at ambient temperature following a standard protocol in which participants had refrained from eating or drinking for at least 3 hours before testing (NIH Technol Assess Statement, 1994).

Clinic Blood Pressure

A trained technician obtained BP measurements using a random zero sphygmomanometer, standardized for cuff size and position. Measurements were made on 4 separate visits during a 3-week period. At each visit, BP was measured in the nondominant arm in the sitting position after an initial rest period of 5 minutes. BP was measured 4 successive times at 2-minute intervals. The first BP measurement of each visit was discarded and the average of the remaining 3 measurements represented the clinic visit BP. The overall clinic BP was then determined by averaging the mean BPs over 4 visits. Measurements were obtained in this standardized manner at baseline and after 6 months.

Data Analysis

Baseline differences among treatment groups were examined using 1-way analysis of variance (ANOVA) for continuous variables and χ^2 tests for categorical variables. Treatment effects were evaluated using analysis of covariance (ANCOVA), with posttreatment values serving as the dependent variables and pretreatment values, age, gender, and treatment as the predictors. Within each model, planned contrasts were used to compare 1) the 2 treatment groups with controls and 2) WM participants with EX subjects. Separate ANCOVA models were used to determine the mediating effects of VO_2 peak and weight loss in predicting improvement in depressive symptoms. Following the intent-to-treat principle, participants were analyzed regardless of adherence to the study protocol.

Results

One hundred thirty three participants were initially randomized for participation (54 in EX, 55 in WM, and 24 in WLC) (Figure 1). Twenty one participants dropped out of the study prior to completion (10 in EX, 9 in WM, and 2 in WLC), nine of whom returned for follow-up assessments. Dropouts did not differ from completers in age, gender, or severity of depressive symptoms, although there was a higher percentage of Caucasian participants among completers ($P < .01$). Participants in the EX group had lower SBP compared with their WM counterparts (SBP = 138.0 vs. 143.6, $p = .024$). No other group differences were observed for any clinical or demographic variables.

Adherence

Participants in the active treatment groups attended an average of 77 exercise sessions (mean sessions = 77.5 (SD = 27.2)) and were exercising at or above their target heart range during approximately 83% of their heart rate checks. Among participants completing the study, participants in the EX group attended approximately 74.8% of exercise sessions and exercised at or above their target heart rate during approximately 84.6% of their sessions. Participants in the WM group attended an average of 78.7% of exercise sessions and exercised at or above their target heart rate during approximately 81.2% of their sessions. There were no significant differences between the EX and WM groups in attendance or adherence to the exercise protocol, although younger participants tended to have lower attendance ($p = .06$). Adherence to the exercise protocol was not associated with pretreatment levels of depressive symptoms ($p = .411$).

Changes in Blood Pressure, Body Weight, and Aerobic Capacity

EX and WM groups showed significant reductions in systolic and diastolic blood pressures at the completion of the study compared with control participants ($F_{1,127} = 10.88, P < .001$; $F_{1,127} = 29.46, p < .001$), although there were no significant differences between the EX and WM groups ($F_{1,127} = 1.95, P = .165$; $F_{1,127} = 1.72, P < .192$). There were significant differences between the 3 groups in weight loss such that participants in the active treatment groups

demonstrated significantly greater weight loss compared with control participants ($F_{1,114} = 30.44, P < .001$) and WM participants showed greater weight loss compared with their EX counterparts ($F_{1,114} = 48.68, P < .001$). Participants in the EX and WM participants groups lost an average of 1.8 and 7.9 kilograms (kgs) while control participants gained an average of .71 kgs.

Participants in the active treatment groups also showed greater improvement in exercise treadmill time (ETT) ($F_{1,115} = 50.83, P < .001$) and peak oxygen consumption (VO_2) ($F_{1,106} = 35.35, P < .001$) compared with controls. In addition, WM participants experienced greater improvements in ETT compared with EX participants ($F_{1,115} = 14.68, P < .01$), although the groups did not differ significantly in VO_2 improvement ($F_{1,106} = 1.08, P = .30$). Analysis of pre-to-post treatment mean differences for treadmill time showed that EX and WM participants increased their ETT by 77 and 136 seconds, respectively, while WLC participants showed a 8 second decline. Similarly, pre-to-post treatment mean gains for VO_2 peak were .29 mL/kgs/min for EX and WM participants, whereas WLC participants exhibited a $-.05$ mL/kgs/min decline.

Changes in depression

Participants exhibited low levels of depression at baseline (mean BDI = 4.8 (SD = 4.3)) and only 21 participants (16%) reported at least “mild” levels of depression[39]. Planned contrasts examining treatment effects on posttreatment depression showed no differences between the EX and WM groups compared with the WLC group ($F_{1,112} = 0.92, p = .339$). Similarly, the EX and WM groups did not differ significantly ($F_{1,112} = 0.08, P = .773$) in post-treatment levels of depression.

Further analysis of treatment differences in depression revealed a significant treatment group by baseline depression severity interaction ($F_{2,106} = 5.75, P < .01$) in predicting posttreatment levels of depression, such that participants in the WM and EX groups with higher BDI scores at baseline demonstrated significantly greater reductions in BDI scores compared with controls. The reduction in depressive symptoms among participants in the active treatment groups was more pronounced in the WM group, although EX participants also improved significantly (Figure 2). In order to examine the magnitude of this effect, we examined the number of participants who exhibited a $> 25\%$ reduction in BDI scores, consistent with previous clinical investigations. Excluding those participants who reported no depressive symptoms at baseline, and could therefore not improve ($n = 19$), approximately 35.4% of EX participants and 38.1% of WM participants showed significant reductions in depressive symptoms, compared to only 21.7% of WLC participants.

Mediators of Change in Depressive Symptoms

In order to investigate potential mediators to the improvement in depressive symptoms, the relationships between improvement in VO_2 peak, weight loss, and depression were examined. Analysis of changes in VO_2 revealed a significant three-way interaction between baseline depression severity, group assignment, and change in VO_2 ($F_{2,92} = 4.64, P = .01$) such that posttreatment levels of depression were related to increased aerobic fitness among participants in the active treatment groups with elevated BDI scores at baseline. An analysis of raw change scores in depression and aerobic capacity among this group of participants showed a significant association between improvement in depression and increased VO_2 peak ($r = .36, P = .02$), indicating that reduced levels of depression were mediated by increases in aerobic capacity. Neither weight loss, nor blood, pressure was associated with reduced levels of depression.

Discussion

Results from the present analysis demonstrate that for men and women with high blood pressure and mild to moderate levels of depression, depressive symptoms may be reduced to a greater extent by an exercise program, either alone or combined with a weight management intervention, compared with participants receiving no treatment. Furthermore, among participants in the active treatment groups, reduced levels of depression were associated with increased aerobic capacity such that participants with greater improvements in VO₂ peak showed greater reductions in depression.

Our results are consistent with previous studies, which indicated that exercise can reduce depressive symptoms in healthy individuals [15,16,40–42] as well as patients with conditions such as heart disease [18–25,43] and arthritis [44]. Because both hypertension [28,29] and elevated depressive symptoms [45,46] are independent risk factors for CHD, an exercise intervention that reduces blood pressure and depression may be especially beneficial among hypertensive patients, a group at elevated risk for the development of CHD.

We observed that the antidepressant effect of exercise was moderated by pretreatment levels of depression. Participants with initially elevated levels of depressive symptoms benefited from exercise, whereas exercise resulted in minimal changes in depression for participants who were initially non-depressed. It is possible that there was a “floor effect” such that participants with low depressive symptoms had no room for improvement, hence only those participants with elevated depressive scores could improve. Alternatively, it is possible that, among individuals with elevated levels of depressive symptoms at baseline, those whose depressive symptoms were more rapidly reduced were able to engage in the exercise portion of the intervention more effectively.

Exercise has repeatedly been shown to reduce depressive symptoms in a variety of depressed populations [7] and to be equally effective [11] compared to anti-depressant therapy. In one of the first clinical trials to evaluate the use of aerobic exercise as a treatment for major depressive disorder (MDD), Blumenthal and colleagues [11] found that a 16-week aerobic exercise intervention was equally effective to treatment with Sertraline in alleviating depressive symptoms. Moreover, in a follow-up analysis from this original trial, Babyak and colleagues [12] found individuals who participated in the exercise portion of the study were less likely to have a recurrent depressive episode 10 months following study completion and that reduced risk was associated with higher levels of physical activity in this subset of patients. Similarly, in a recent meta-analysis of 11 treatment outcome studies utilizing physical activity in the treatment of depression, Stathopoulou and colleagues [7] found a clear advantage of exercise compared to control conditions.

Mechanisms by which exercise may reduce depressive symptoms by affecting changes in central monoamines, increasing hypothalamic-pituitary-adrenal (HPA) axis regulation, altering endorphin levels, increasing positive self-evaluations, and by enhancing cardiopulmonary functioning. Improved aerobic capacity assessed by VO₂ peak has been associated with improvement in depression in several previous studies among both cardiac patients [18,19,47] and healthy patients [48,49]. Recent studies have shown that decreased aerobic capacity predicts increased levels of depressive symptoms as early as two weeks following exercise withdrawal among healthy individuals [50]. Among individuals with depression, Dunn and colleagues (CIT) recently demonstrated that a dose response relationship may exist between increased levels of aerobic fitness and decreased levels of depression. Our results provide further evidence to support this relationship by showing that training-induced improvement in aerobic capacity was associated with a reduction in symptoms of depression. These findings suggest that improved fitness may have contributed to the antidepressant effects

of the exercise interventions. In contrast to previous findings [51], weight reduction was not significantly associated with improvement in depression.

The findings of this study had several notable limitations. First, because participants exhibited relatively low levels of depression at baseline our findings may not be generalizable to individuals with more severe depressive symptoms. Second, because depressive symptoms were assessed currently at two time points, the direction of the relationship was between changes in depressive symptoms and changes in aerobic fitness is unclear: improved aerobic capacity could result in reduced depression, or reduced depression may contribute to improved aerobic capacity. Finally, the WLC group in our study did not receive as much attention from the study staff as the active treatment groups, which may have affected their mood over the course of the 6-month treatment program. Future studies investigating this relationship would be improved with the addition of an attentional control group.

Results of this study may have important clinical implications. Previous studies have shown that depression is associated with adverse medical outcomes [52]. Because depressive symptoms have been associated with a greater incidence of adverse events in the presence of hypertension [33,53], we feel that our findings may have important implications for the management of depression among hypertensive individuals. Furthermore, given that depression has been associated with a greater risk for both the development of CHD [54] and increased mortality following cardiac events [55], interventions that alleviate depressive symptoms in this at-risk population may have important public health implications. Future research should examine the benefits of exercise on clinical outcomes among depressed hypertensive patients, including assessment of other important biomarkers of risk.

Acknowledgements

This study was supported by grants HL 49572, HL 59672, HL 65503, HC 55142, MH 49679, and HL 59672 from the National Institutes of Health and grant M01-RR-30 from the General Clinical Research Center Program, National Center for Research Resources, National Institutes of Health.

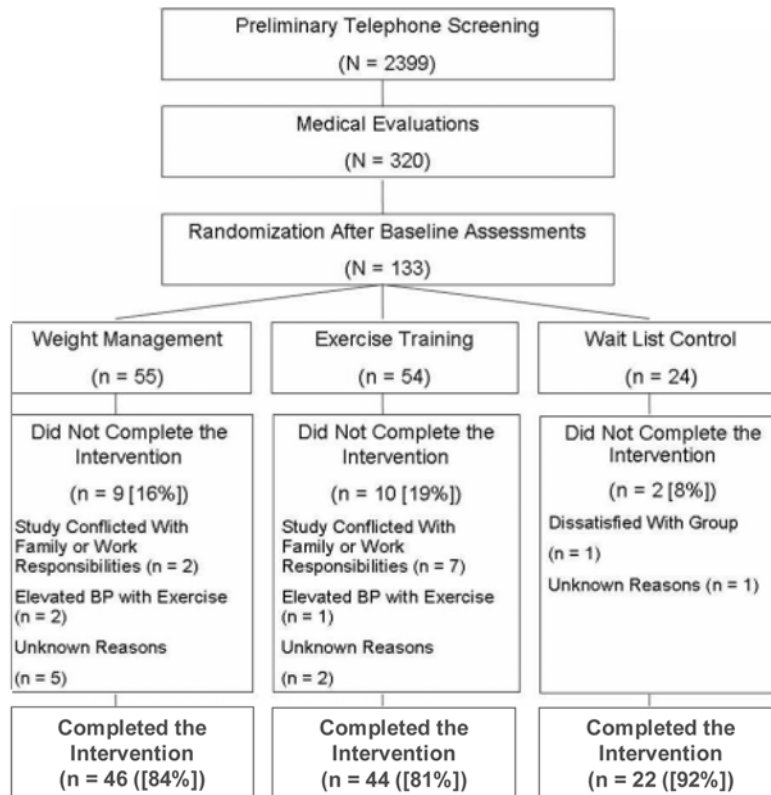
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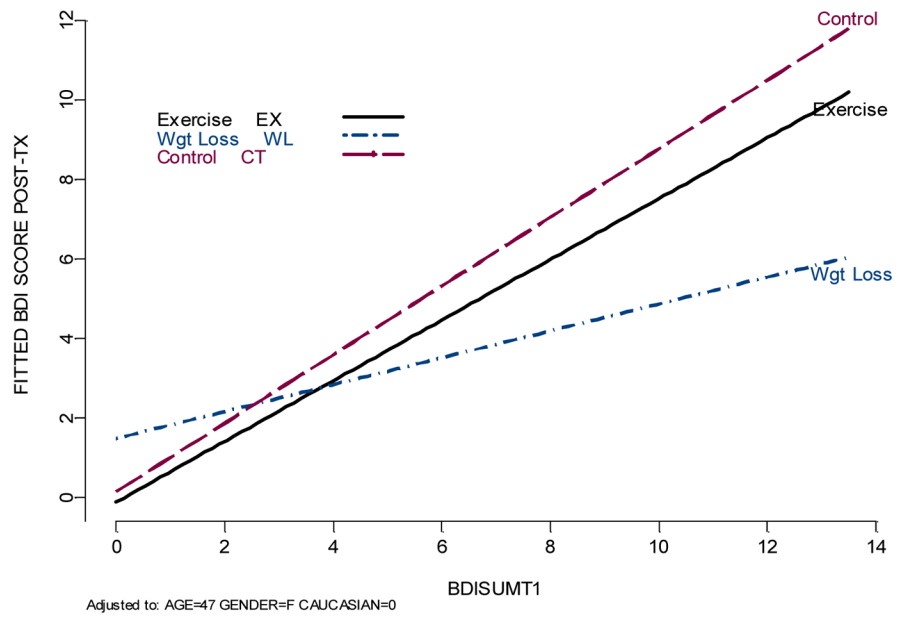
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1. .



2. .

Table 1

Background characteristics*

| Variable | EX (n = 54) | WM (n = 55) | WLC (n = 24) | Total (n = 133) |
|-----------------------------|--------------|--------------|--------------|-----------------|
| Age, mean (SD), y | 46.6 (8.2) | 48.5 (9.9) | 47.2 (7.4) | 47.5 (8.8) |
| Males | 25 (46.3) | 21 (38.2) | 13 (54.2) | 59 (44.4) |
| Caucasians | 41 (75.9) | 44 (80.0) | 15 (62.5) | 100 (75.2) |
| College Degree | 38 (70.4) | 34 (61.8) | 15 (62.5) | 87 (65.4) |
| SBP, mean (SD), mm Hg | 141.1 (14.8) | 145.5 (16.3) | 145.7 (14.0) | 143.7 (15.3) |
| DBP, mean (SD), mm Hg | 91.0 (8.1) | 90.6 (7.6) | 91.5 (7.1) | 90.9 (7.7) |
| Body weight, mean (SD), KGS | 95.5 (14.5) | 93.4 (16.9) | 93.1 (16.9) | 94.2 (15.9) |
| BDI, mean (SD) | 5.2 (4.1) | 4.0 (4.1) | 5.7 (5.0) | 4.8 (4.5) |

* Data are given as number (percentage) unless otherwise indicated. SBP indicates systolic blood pressure; DBP indicates diastolic blood pressure; mm Hg indicates millimeters of mercury; KGS indicates kilograms; BDI indicates Beck Depression Inventory.