

Articles are selected for CME credit designation on the basis of our assessment of the needs of readers of *The Primary Care Companion*, with the purpose of providing readers with a curriculum of CME articles on a variety of topics throughout each volume. There are no prerequisites for participation in this CME activity.

To obtain credit, please study the designated article and complete the Posttest.

Accreditation Statement

Physicians Postgraduate Press, Inc. is accredited by the Accreditation Council for Continuing Medical Education to provide continuing medical education for physicians.

Credit Designation

Physicians Postgraduate Press, Inc. designates this educational activity for up to 1 Category 1 credit toward the American Medical Association Physician's Recognition Award. Each participant should claim only those credits that he/she actually spent in the educational activity.

Date of Original Release/Review

This educational activity is eligible for CME credit through June 30, 2006. The latest review of this material was May 2004.

Educational Objective

After studying the article by Craft and Perna, you will be able to:

• Counsel patients on the adoption of exercise as an adjunct treatment for depression.

This pretest is designed to facilitate your study of the material.

- **1.** Aerobic and nonaerobic forms of exercise *do not* appear to be equally effective in alleviating symptoms of clinical depression.
 - a. True
 - b. False

Pretest answer and Posttest on page 112.

Disclosure of Off-Label Usage

The authors have determined that, to the best of their knowledge, no investigational information about pharmaceutical agents has been presented in this article that is outside U.S. Food and Drug Administration–approved labeling.

 Answers to Vol. 5, No. 1 Posttest 2003

 1. c
 2. a
 3. d
 4. b
 5. d



The Benefits of Exercise for the Clinically Depressed

Lynette L. Craft, Ph.D., and Frank M. Perna, Ed.D., Ph.D.

Millions of Americans suffer from clinical depression each year. Most depressed patients first seek treatment from their primary care providers. Generally, depressed patients treated in primary care settings receive pharmacologic therapy alone. There is evidence to suggest that the addition of cognitive-behavioral therapies, specifically exercise, can improve treatment outcomes for many patients. Exercise is a behavioral intervention that has shown great promise in alleviating symptoms of depression. The current review discusses the growing body of research examining the exercise-depression relationship that supports the efficacy of exercise as an adjunct treatment. Databases searched were Medline, PsycLit, PubMed, and SportsDiscus from the years 1996 through 2003. Terms used in the search were clinical depression, depression, exercise, and physical activity. Further, because primary care physicians deliver important mental health services to the majority of depressed patients, several specific recommendations are made regarding counseling these patients on the adoption and maintenance of exercise programs. (Prim Care Companion J Clin Psychiatry 2004;6:104-111)

Received Dec. 2, 2003; accepted Feb. 5, 2004. From the Division of Psychiatry, Boston University School of Medicine, Boston, Mass. This manuscript was partially supported by the National Institutes

In the spirit of full disclosure and in compliance with all ACCME Essential Areas and Policies, the faculty for this CME activity were asked to complete a full disclosure statement. The information received is as follows: Drs. Craft and Perna have no significant commercial relationships to disclose relative to the presentation.

Corresponding author and reprints: Lynette L. Craft, Ph.D., Division of Psychiatry, Boston University School of Medicine, 85 E. Newton Street, Room M-619, Boston, MA 02118 (e-mail: craftlyn@bu.edu).

Depression affects roughly 9.5% of the U.S. adult population each year, and it is estimated that approximately 17% of the U.S. population will suffer from a major depressive episode at some point in their lifetime.^{1,2} Depression has been ranked as the leading cause of disability in the United States, with over \$40 billion being spent each year on lost work productivity and medical treatment related to this illness.³⁻⁶ Recent research suggests that between the years of 1987 and 1997, the rate of outpatient treatment for depression in the United States tripled and that health care costs related to this disorder continue to rise.⁷

The vast majority of those suffering from depression first seek treatment from their primary care providers.⁸ As such, it has been estimated that the prevalence of depression in primary care settings is 3 times that in community samples, and rates for minor depression and dysthymia are even greater, ranging from 5% to 16% of patients.^{8,9} Depressed patients treated in primary care settings receive predominantly pharmacologic therapy, with fewer receiving adjunct cognitive or behavioral interventions.⁷ As a result, it is likely that many of these patients are not educated regarding nonpharmacologic strategies for managing the symptoms of their depression. Treatment of clinical depression can be improved by the addition of cognitive-behavioral therapies,10 and by exercise. Research has also shown that depressed patients are less fit and have diminished physical work capacity on the order of 80% to 90% of age-predicted norms,¹¹⁻¹⁴ which in turn may contribute to other physical health problems. Therefore, primary care providers are uniquely positioned to promote behavioral approaches, such as exercise, that complement pharmacologic treatment and may ultimately provide relief from this chronic and often treatmentresistant disorder as well as enhance overall physical well-being. The current review discusses the growing body of research examining the exercise-depression relationship. Databases searched were Medline, PsycLit, PubMed, and SportsDiscus from the years 1996 through 2003. Terms used in the search were *clinical depression*, depression, exercise, and physical activity.

EXERCISE AND CLINICAL DEPRESSION

Involvement in structured exercise has shown promise in alleviating symptoms of clinical depression. Since the early 1900s, researchers have been interested in the association between exercise and depression. Early case studies concluded that, at least for some, moderate-intensity exercise should be beneficial for depression and result in a happier mood.^{15,16} Further, a relationship between physical work capacity (PWC) and depression appeared to exist,^{11–14} but the directional nature of this relationship could not be addressed via case and cross-sectional studies. However, researchers have remained interested in the antidepressant effects of exercise and more recently have utilized experimental designs to study this association.

104

of Health General Clinical Research Center Grant #M01RR005333 to Boston University School of Medicine.

Many studies have examined the efficacy of exercise to reduce symptoms of depression, and the overwhelming majority of these studies have described a positive benefit associated with exercise involvement. For example, 30 community-dwelling moderately depressed men and women were randomly assigned to an exercise intervention group, a social support group, or a wait-list control group.¹⁷ The exercise intervention consisted of walking 20 to 40 minutes 3 times per week for 6 weeks. The authors reported that the exercise program alleviated overall symptoms of depression and was more effective than the other 2 groups in reducing somatic symptoms of depression (reduction of 2.4 [walking] vs. 0.9 [social support] and 0.4 [control] on the Beck Depression Inventory [BDI], p < .05). Doyne et al.¹⁸ utilized a multiple baseline design to evaluate the effectiveness of interval training in alleviating symptoms of depression. The participants exercised on a cycle ergometer 4 times per week, 30 minutes per session, for 6 weeks. This treatment was compared with an attention-placebo control condition in which subjects listened to audiotapes of "white noise" that they were told was subliminal assertiveness training. Results indicated that the aerobic training program was associated with a clear reduction in depression compared with the control condition, and the improvements in depression were maintained at 3 months post intervention (BDI mean reduction of 14.4 points from baseline, p < .05). In another study, just 30 minutes of treadmill walking for 10 consecutive days was sufficient to produce a clinically relevant and statistically significant reduction in depression (reduction of 6.5 points from baseline on the Hamilton Rating Scale for Depression [HAM-D], p < .01).¹⁹

Research also suggests that the benefits of exercise involvement may be long lasting.²⁰ Depressed adults who took part in a fitness program displayed significantly greater improvements in depression, anxiety, and self-concept than those in a control group after 12 weeks of training (BDI reduction of 5.1 [fitness program] vs. 0.9 [control], p < .001). The exercise participants also maintained many of these gains through the 12-month follow-up period.²⁰

While most studies have employed walking or jogging programs of varying lengths, the efficacy of nonaerobic exercise has also been assessed. For example, in comparison with a control condition, resistance-training programs reduced symptoms of depression (resistance training vs. control resulted in BDI reduction of 11.5 vs. 4.6, respectively, p < .01, and HAM-D reduction of 7.0 vs. 2.5, respectively, p < .01).²¹ Aerobic and nonaerobic modes of exercise have also been compared to determine if certain types of activities are more effective than others. Doyne and colleagues²² compared the efficacy of running with that of weight lifting. Forty depressed women served as participants and were randomly assigned to running, weight lifting, or a wait-list control group. Participants

were asked to complete 4 training sessions each week for the 8 weeks of the program. Depression was assessed at mid- and post-treatment and at 1, 7, and 12 months followup. Results indicated that the 2 activities were not significantly different, and that both types of exercise were sufficient to reduce symptoms of depression (running vs. weights vs. control resulted in BDI reduction of 11.1 vs. 13.6 vs. 0.8, respectively, p < .01, and HAM-D reduction of 6.7 vs. 8.7 vs. a 1.0 increase, respectively, p < .01). Further, there were no differences between the 2 treatment groups during follow-up with respect to the percentage of participants who remained nondepressed. Similarly, a study by Martinsen et al.²³ assessed 90 depressed inpatients who were randomly assigned to aerobic or nonaerobic exercise. Aerobic exercise consisted of jogging or brisk walking, and nonaerobic exercise included strength training, relaxation, coordination, and flexibility training. The program was 8 weeks in length, and participants exercised for 60 minutes, 3 times per week. Those in the aerobic group exhibited an increase in PWC compared with those in the nonaerobic group. However, both groups experienced a significant reduction in depression score (p < .001), but there were no significant differences between the groups with respect to the magnitude of change in depression score (p > .10).

Additionally, exercise compares quite favorably with standard care approaches to depression in the few studies that have evaluated their relative efficacy. For example, running has been compared with psychotherapy in the treatment of depression, with results indicating that running is just as effective as psychotherapy in alleviating symptoms of depression (Symptom Checklist-Depression reduction in mean item score of 1.9 [running] vs. 1.6 [therapy], NS).²⁴ The benefits of running have been compared with cognitive therapy alone and a combination of running and cognitive therapy.²⁵ Participants in this study were randomly assigned to running only, running and therapy, or cognitive therapy only. The treatment was 10 weeks in length. The running group met 3 times per week and exercised for 20 minutes per session. Those in the therapy-only group met with a therapist for 60 minutes once a week. Those in the combination group received 10 individual sessions with a therapist and also ran 3 times per week. There were no significant differences between these 3 groups, with all groups displaying a significant reduction in depression, and the positive benefits were still present at the 4-month follow-up (BDI reduction of 10.9 [running] vs. 11.0 [therapy] vs. 7.7 [combined]; main effect for time, p < .001; group × time interaction, NS, p > .05).

The efficacy of exercise relative to psychotropic medication has also been investigated. Blumenthal and colleagues²⁶ randomly assigned 156 moderately depressed men and women to an exercise, medication, or exercise and medication group. Those in the exercise group walked

or jogged on a treadmill at 70% to 85% of heart rate reserve for 30 minutes 3 times per week for 16 weeks. Those in the medication group received sertraline, and a psychiatrist evaluated medication efficacy, assessed side effects, and adjusted dosages accordingly at 2, 6, 10, 14, and 16 weeks. Those in the combination group received both medication and exercise prescription according to the procedures described previously. Results showed that while medication worked more quickly to reduce symptoms of depression, there were no significant differences among treatment groups at 16 weeks (HAM-D: F = 0.96, df = 2,153; p = .39; BDI: F = 0.90, df = 2,153; p = .40).The percentage of patients in remission from their depression at 16 weeks did not differ among groups (60.4% [exercise] vs. 68.8% [medication] vs. 65.5% [combination], p = .67). Therefore, exercise was as effective as medication for reducing symptoms of depression in that sample. Interestingly, 10-month follow-up of those participants revealed that exercise group members (70%) had significantly (p = .028) lower rates of depression than those in the medication (48%) or the combination groups (54%).²⁷ Finally, at 10 months, regular exercise involvement was a significant predictor of lower rates of depression (OR = 0.49, CI = 0.32 to 0.74, p < .01).²⁶

META-ANALYTIC FINDINGS

While relatively few studies have been described here for illustrative purposes, there are now a large number of studies that support the efficacy of exercise in reducing symptoms of depression. Further, while the early research in this area suffered from a variety of methodological limitations (e.g., small samples, lack of random assignment, lack of control groups), current researchers have addressed these design issues, and presently there are multiple studies that have utilized experimental designs or employed a randomized clinical trial approach. Metaanalysis provides one means of summarizing this growing body of primary research and identifying variables that may moderate the effect of exercise on depression. Effect sizes (ESs) are calculated for each study and weighted to correct for positive bias that can result from small sample sizes. The statistical analyses of these ESs allow the researcher to investigate study and subject characteristics that may moderate the exercise-depression relationship as well as compare subsets of samples from the original studies and make comparisons (e.g., male vs. female, mildly vs. moderately depressed) that were not directly made within each of the individual primary research articles.

There have been several meta-analyses conducted on the literature examining the relationship between exercise and depression. North et al.²⁸ included 80 studies in their meta-analysis and reported an overall mean ES of –0.53, indicating that exercise reduced depression scores by approximately one half a standard deviation compared with those in comparison groups. That meta-analysis included primary research studies that had examined the exercisedepression relationship in a variety of samples (e.g., college students, medical patients, depressed and mentally ill patients, and normal adults). When those authors analyzed the subgroup of studies that had utilized clinical populations (e.g., substance abusers, post–myocardial infarction patients, hemodialysis patients), they reported an even larger ES of -0.94. However, patients suffering from depression secondary to a medical condition may be qualitatively different from those who suffer from clinical depression as the primary illness. As such, exercise interventions that are effective for medical populations may not be as effective in treating depression when it is the primary disorder.²⁹

Craft and Landers³⁰ attempted to further clarify this relationship and included only studies in which individuals diagnosed with clinical depression served as participants. Thirty studies were included in that meta-analysis, and an overall mean ES of -0.72 was reported. Several variables related to study quality, subject characteristics, and exercise program characteristics were coded and examined in an attempt to determine potential moderators of this relationship. Interestingly, exercise program characteristics such as duration, intensity, frequency, and mode of exercise did not moderate the effect. In fact, only the length of the exercise program was a significant moderator, with programs 9 weeks or longer being associated with larger reductions in depression. Further, exercise was equally effective across a variety of patient subgroups. For example, subject characteristics such as age, gender, and severity of depression did not emerge as significant moderators. Finally, when compared with other traditional treatments for depression, exercise was just as beneficial and not significantly different from psychotherapy, pharmacologic therapy, and other behavioral interventions.³⁰

Lawlor and Hopker³¹ more recently conducted a metaanalysis that included only randomized controlled trials of clinically depressed patients. Fourteen studies were included in their meta-analysis. They reported an overall mean ES of -1.1 for exercise interventions compared with no-treatment control groups. Exercise was also as effective as cognitive therapy in alleviating symptoms of depression, with a nonsignificant ES of -0.3 emerging.

Analyses of subsamples within these meta-analyses indicate that exercise is effective for both men and women of all ages as well as those who are initially more severely depressed. Further, comparison of exercise program characteristics across the studies yielded nonsignificant ESs, indicating that exercise need not be lengthy or intense and that fitness gains are not necessary for patients to experience positive benefits. Therefore, in light of this additional meta-analytic support, researchers have begun to recommend that depressed individuals should adopt physically active lifestyles to help manage their symp-

udy Characteristic	No. of Studies
mple size, N	
< 30	21
30–60	10
> 60	6
bject assignment	
Stratified/block random	8
Random	19
Intact groups	1
Single group (pre/post)	9
epression measure ^b	
Beck Depression Inventory	25
Zung Self-Rating Depression Scale	6
Symptom Checklist-90	6
Hamilton Rating Scale for Depression	4
Brink Geriatric Scale	2
Children's Depression Scale	2
Clinical Interview Schedule	1
DSM-IV and others	2
pe of intervention	
Running	8
Running and walking	16
Walking	2
Other aerobic	6
Nonaerobic	4
Not reported	1
nder	
Men	3
Women	5
Mixed	29

Table 1. Characteristics of 37 Studies Included in Exercise			
and Depression Meta-Analyses ^a			

toms.^{30,32,33} Table 1 provides basic characteristics of the studies utilizing clinical populations that were included in the 3 meta-analyses discussed above.^{28,30,31} Readers are referred to each of the meta-analyses for additional information regarding individual primary research articles. Further, a review by Dunn and colleagues³⁴ includes a summary of study and participant characteristics for quasiexperimental and experimental designs.

The data from the 3 meta-analyses discussed previously^{28,30,31} resulted in overall ESs of -0.72, -0.94, and -1.1. Converting these ESs to a binomial effect size display allows one to examine the practical clinical significance of these effect sizes.³⁵ These values reflect an increase in success rate due to treatment (i.e., exercise) of 67%, 71%, and 74%, respectively, and such promising treatment outcomes are notable. In medical settings, clinical guidelines suggest that a 50% reduction in symptoms during the treatment phase is considered a treatment response. Researchers have argued that this criterion is more clinically relevant than statistically significant results, which can occur even if the participants have not experienced a 50% reduction in the symptoms of their depression.³⁴ Therefore, even when using this more clinically relevant guideline, the meta-analytic data provide support for the efficacy of exercise in reducing symptoms

of depression. While this discussion is not meant to argue against the use of antidepressant medication or psychological therapies, there is strong evidence to advocate the use of exercise as a potentially powerful adjunct to existing treatments.

PROPOSED MECHANISMS FOR THE EXERCISE-DEPRESSION RELATIONSHIP

While the research is consistent and points to a relationship between exercise and depression, the mechanisms underlying the antidepressant effects of exercise remain unclear. Several credible physiologic and psychological mechanisms have been described, such as the thermogenic hypothesis,³⁶ the endorphin hypothesis,^{37,38} the monoamine hypothesis,^{39–42} the distraction hypothesis,⁴³ and the enhancement of self-efficacy.^{29,43-46} However, there is little research evidence to either support or refute most of these theories.

Thermogenic Hypothesis

The thermogenic hypothesis suggests that a rise in core body temperature following exercise is responsible for the reduction in symptoms of depression. DeVries³⁶ explains that increases in temperature of specific brain regions, such as the brain stem, can lead to an overall feeling of relaxation and reduction in muscular tension. While this idea of increased body temperature has been proposed as a mechanism for the relationship between exercise and depression, the research conducted on the thermogenic hypothesis has examined the effect of exercise only on feelings of anxiety rather than depression.^{36,47,48}

Endorphin Hypothesis

The endorphin hypothesis predicts that exercise has a positive effect on depression due to an increased release of β-endorphins following exercise. Endorphins are related to a positive mood and an overall enhanced sense of well-being. This line of research has not been without criticism. The debate remains as to whether plasma endorphins reflect endorphin activity in the brain. Some^{37,38} have argued that even if peripheral endorphin levels are not reflective of brain chemistry, they could still be associated with a change in mood or feelings of depression. Several studies have shown increases in plasma endorphins following acute and chronic exercise⁴⁹⁻⁵¹; yet, it remains unclear if these elevations in plasma endorphins are directly linked to a reduction in depression. Lastly, the phenomenon of runner's high, often attributed to endorphin release, is not blocked by naloxone injection, an opiate antagonist.52,53

Monoamine Hypothesis

The monoamine hypothesis appears to be the most promising of the proposed physiologic mechanisms. This

hypothesis states that exercise leads to an increase in the availability of brain neurotransmitters (e.g., serotonin, dopamine, and norepinephrine) that are diminished with depression. These neurotransmitters increase in plasma and urine following exercise, but whether exercise leads to an increase in neurotransmitters in the brain remains unknown.^{40–42} Animal studies suggest that exercise increases serotonin and norepinephrine in various brain regions,^{39,54–56} but, to date, this relationship has not been studied in humans.

Therefore, while several physiologic mechanisms remain plausible, methodological difficulties have prevented this line of research from advancing. Martinsen⁵⁷ discusses how testing biochemical hypotheses is often difficult in humans due to the invasive procedures necessary to obtain samples (e.g., spinal taps for cerebrospinal fluid samples). Further, biochemical samples obtained from blood or other bodily fluids may not directly reflect the activity of these compounds in the brain.³⁹ Hopefully, with the advent of new less invasive neuroimaging techniques, future researchers can examine whether exercise leads to the neurochemical changes in the brain predicted by these physiologic hypotheses.

Distraction Hypothesis

Several psychological mechanisms have also been proposed. As was the case with the physiologic mechanisms, many of these theories have not been tested extensively. The distraction hypothesis suggests that physical activity serves as a distraction from worries and depressing thoughts.⁴³ In general, the use of distracting activities as a means of coping with depression has been shown to have a more positive influence on the management of depression and to result in a greater reduction in depression than the use of more self-focused or introspective activities such as journal keeping or identifying positive and negative adjectives that describe one's current mood.^{58,59}

Exercise has been compared with other distracting activities such as relaxation, assertiveness training, health education, and social contact.^{17,18,23,60} Results have been inconclusive, with exercise being more effective than some activities and similar to others in its ability to aid in the reduction of depression. However, exercise is known to increase positive affect, which is diminished in depressed patients and is not augmented by distraction activities. The diminished capacity to experience positive affect is an essential distinguishing symptom in clinical depression.

Self-Efficacy Hypothesis

The enhancement of self-efficacy through exercise involvement may be another way in which exercise exerts its antidepressant effects. Self-efficacy refers to the belief that one possesses the necessary skills to complete a task as well as the confidence that the task can actually be completed with the desired outcome obtained. Bandura³⁶ describes how depressed people often feel inefficacious to bring about positive desired outcomes in their lives and have low efficacy to cope with the symptoms of their depression. This can lead to negative self-evaluation, negative ruminations, and faulty styles of thinking. It has been suggested that exercise may provide an effective mode through which self-efficacy can be enhanced based on its ability to provide the individual with a meaningful mastery experience. Research examining the association between physical activity and self-efficacy in the general population has focused predominantly on the enhancement of physical self-efficacy and efficacy to regulate exercise behaviors. The relationship between exercise and self-efficacy in the clinically depressed has received far less attention. The findings of the few studies that have examined this relationship have been equivocal as to whether exercise leads to an enhancement of generalized feelings of efficacy.^{23,61} However, 1 recent study⁶² has reported that involvement in an exercise program was associated with enhanced feelings of coping self-efficacy, which, in turn, were inversely related to feelings of depression.

More research is needed to determine which, if any, of the mechanisms described herein are important moderators of the exercise effect. It is highly likely that a combination of biological, psychological, and sociological factors influence the relationship between exercise and depression. This is consistent with current treatment for depression in which the effects of pharmacotherapy and psychotherapy on depression are additive and address biological, psychological, and sociological aspects of the patient. There may also be individual variation in the mechanisms or combination of mechanisms mediating this relationship.³² Additionally, different mechanisms may be important at specific times in the natural course of depression. Until more is known about the possible mechanisms, this relationship may be best studied utilizing a biopsychosocial approach.

PRACTICAL CONSIDERATIONS

With increasing evidence to support the efficacy of this behavioral intervention in reducing symptoms of depression, we encourage primary care providers to recommend exercise involvement to their depressed patients. However, if physicians are to recommend physical activity as adjunct therapy, there are a few practical considerations that should be addressed. First, patients with depression are typically sedentary and may lack motivation to begin an exercise program. The American College of Sports Medicine⁶³ has made the recommendation that all adults should exercise at least 30 minutes per day, on all or most (5) days per week, at a moderate-vigorous intensity. Such a recommendation is likely to be overwhelming to some-

one who is currently sedentary and depressed. Based on the meta-analytic findings in this area, an exercise prescription of 20 minutes per day, 3 times per week, at a moderate intensity is sufficient to significantly reduce symptoms of depression.³⁰ Therefore, we suggest that physicians begin with this prescription as an initial recommendation.

Physicians should counsel patients to begin slowly and choose an enjoyable mode of exercise. For many people, walking is often a cost-effective option, and the physician should ask patients to identify a safe place to walk and a place to walk indoors during inclement weather (e.g., a shopping mall). The primary care provider may want to encourage the patient to first work on the frequency of his or her exercise activities, that is, setting a goal of 3 exercise sessions per week, at a comfortable pace, for a manageable length of time (e.g., 10 minutes). Once the patient becomes confident that the exercise frequency is under his or her control, the duration of the exercise bout can gradually be increased.

The final exercise characteristic that should be addressed is the exercise intensity. Reminding patients that exercise need not be lengthy or intense to enhance mood is also a good strategy. For patients with a low level of fitness, which characterizes most depressed patients, moderate-intensity exercise (60%–80% maximum heart rate) generally produces greater enjoyment than more intense activity.^{64–69} Further, in the general population, beginning a moderate-intensity exercise program is associated with roughly one half the dropout rate of beginning a vigorous-intensity exercise program.⁷⁰ Therefore, suggesting that patients begin with lower-intensity exercise activities that they find enjoyable should result in a more positive exercise experience for the patient and increase the likelihood of maintained exercise involvement.

Some data suggest that exercise may also improve sleep quality for some patients.^{71,72} However, at this time, causal inferences cannot be made, and specific sleep data for depressed patients are needed. Results are equivocal as to whether time of day of exercise plays a role in this relationship. As such, patients should be encouraged to exercise at a time that is most convenient for them. For some patients, depressive symptoms worsen later in the day and exercise may be easier to schedule and manage if conducted in the morning or afternoon. With limited data to support the notion that early evening exercise facilitates better sleep, physicians are encouraged to counsel patients that sleep quality may be improved by exercise, and patients should exercise when it is most convenient or feels best for them.

Briefly instructing patients on the benefits of using self-monitoring techniques (exercise logs, pedometers) and setting small, achievable goals can also be significant points of discussion when patients are adopting exercise programs. In the general population, research has shown that addressing a number of key factors specific to the individual (e.g., exercise self-efficacy, social support for exercise, and perceived barriers to exercise), as well as continued brief supportive contact with the researcher or physician, can increase the rate of adherence to the exercise program by up to 25%.^{73–76} The use of self-monitoring and brief follow-up contact with the physician is aimed at increasing the patient's awareness of the immediate rewards of exercise, which is more effective in promoting adherence than focusing on distant outcomes. Therefore, primary care physicians are encouraged to regularly inquire (e.g., during office visits) about exercise involvement and reinforce any positive benefits the patient reports experiencing.

SUMMARY

The mechanisms underlying the antidepressant effects of exercise remain in debate; however, the efficacy of exercise in decreasing symptoms of depression has been well established. Data regarding the positive mood effects of exercise involvement, independent of fitness gains, suggest that the focus should be on frequency of exercise rather than duration or intensity until the behavior has been well established. The addition of self-monitoring techniques may increase awareness of the proximal benefits of exercise involvement, which is generally reinforcing to the patient.

Physician advice is likely to go a long way toward providing motivation and support for exercise. Follow-up contact may also be important during exercise adoption. While this follow-up may present a time challenge to the provider, less time-consuming interactions such as brief telephone contact and automated telephone contact have been shown to increase adherence to exercise programs.^{77–80} Finally, it is interesting to note that while depression may be an additional risk factor for exercise noncompliance, reported drop-out rates among depressed patients are not too different from those in the general population.^{13,21–23,26}

Drug names: naloxone (Suboxone, Narcan, and others), sertraline (Zoloft).

REFERENCES

- Kessler RC, McGonagle KA, Zhao S, et al. Lifetime and 12-month prevalence of DSM-III-R psychiatric disorders in the United States: results from the National Comorbidity Survey. Arch Gen Psychiatry 1994;51: 8–19
- National Institute of Mental Health. The numbers count: mental disorders in America. Available at: http://www.nimh.gov. Accessed Sept 3, 2003
- Broadhead WE, Blazer DG, George LK, et al. Depression, disability days, and days lost from work in a prospective epidemiologic survey. JAMA 1990;264:2524–2528
- Murray CJ, Lopez AD. Evidence-based health policy lessons form the Global Burden of Disease Study. Science 1996;274:740–743
- World Health Organization. Depression. Available at http:// www.who.org. Accessed Nov 15, 2003

- Zerihun M. Depression drains workplace productivity. Available at: http://www.depressionnet.org. Accessed Oct 31, 2001
- Olfson M, Marcus SC, Druss B, et al. National trends in the outpatient treatment of depression. JAMA 2002;287:203–209
- Pincus JA, Pechura CM, Elinson L, et al. Depression in primary care: linking clinical and systems strategies. Gen Hosp Psychiatry 2001;23:311–318
- Frank E, Rucci P, Stat D, et al. Correlates of remission in primary care patients treated for minor depression. Gen Hosp Psychiatry 2002;24: 12–19
- Young JE, Weinberger AD, Beck AT. Cognitive therapy for depression. In: Barlow DH, ed. Clinical Handbook of Psychological Disorders, Third Edition. New York, NY:Guilford Press; 2001
- Morgan WP. Selected physiological and psychomotor correlates of depression in psychiatric patients. Res Q 1968;30:1037–1043
- Morgan WP. A pilot investigation of physical working capacity in depressed and nondepressed males. Res Q 1969;40:859–861
- Martinsen EW, Medhus A, Sandvik L. Effects of aerobic exercise on depression: a controlled study [abstract]. Br Med J (Clin Res ED) 1985;291:109
- Martinsen EW, Strand J, Paulsson G, et al. Physical fitness levels in patients with anxiety and depressive disorders. Int J Sports Med 1989; 10:58–61
- Franz SI, Hamilton GV. The effects of exercise upon retardation in conditions of depression. Am J Psychiatry 1905;62:239–256
- Vaux CL. A discussion of physical exercise and recreation. Am J Phys Med 1926;6:303–333
- McNeil JK, LeBlanc EM, Joyner M. The effect of exercise on depressive symptoms in the moderately depressed elderly. Psychol Aging 1991;6:487–488
- Doyne EJ, Chambless DL, Beutler LE. Aerobic exercise as a treatment for depression in women. Behav Ther 1983;14:434–440
- Dimeo F, Baurer M, Varahram I, et al. Benefits from aerobic exercise in patients with major depression: a pilot study. Br J Sports Med 2001;35: 114–117
- DiLorenzo TM, Bargman EP, Stucky-Ropp R, et al. Long-term effects of aerobic exercise on psychological outcomes. Prev Med 1999;28:75–85
- Sing NA, Clements KM, Fiatarone MA. A randomized controlled trial of progressive resistance training in depressed elders. J Gerontol 1997; 52:M27–M35
- Doyne EJ, Ossip-Klein DJ, Bowman ED, et al. Running versus weight lifting in the treatment of depression. J Consult Clin Psychol 1987;55: 748–754
- Martinsen EW, Hoffart A, Solberg O. Comparing aerobic and nonaerobic forms of exercise in the treatment of clinical depression: a randomized trial. Compr Psychiatry 1989;30:324–331
- Greist JH, Klein MH, Eischens RR, et al. Running as treatment for depression. Compr Psychiatry 1979;20:41–54
- Fremont J, Craighead LW. Aerobic exercise and cognitive therapy in the treatment of dysphoric moods. Cognit Ther Res 1987;11:241–251
- Blumenthal JA, Babyak MA, Moore KA, et al. Effects of exercise training on older patients with major depression. Arch Intern Med 1999;159: 2349–2356
- Babyak M, Blumenthal JA, Herman S, et al. Exercise treatment for major depression: maintenance of therapeutic benefit at 10 months. Psychosom Med 2000;62:633–638
- North TC, McCullagh P, Tran ZV. Effects of exercise on depression. Exerc Sport Sci Rev 1990;18:379–415
- Simons AD, McGowan CR, Epstein LH, et al. Exercise as a treatment for depression: an update. Clin Psychol Rev 1985;5:553–568
- Craft LL, Landers DM. The effect of exercise on clinical depression and depression resulting from mental illness: a meta-analysis. J Sport Exerc Psychol 1998;20:339–357
- Lawlor DA, Hopker SW. The effectiveness of exercise as an intervention in the management of depression: systematic review and meta-regression analysis of randomized controlled trials. BMJ 2001;322:763–767
- Fox KR. The influence of physical activity on mental well-being. Public Health Nutr 1999;2:411–418
- Mutrie N. The relationship between physical activity and clinicallydefined depression. In: Biddle SJH, Fox KR, Boutcher SH, eds. Physical Activity and Psychological Well-Being. London, England: Routledge; 2000
- 34. Dunn AL, Trivedi MH, O'Neil HA. Physical activity dose-response

effects on outcomes of depression and anxiety. Med Sci Sports Exerc $2000;33{:}s587{-}s597$

- Rosenthal R. The evaluation of meta-analytic procedures and metaanalytic results. In: Rosenthal R, ed. Meta-Analytic Procedures for Social Research. Newbury Park, Calif: Sage; 1984
- deVries HA. Tranquilizer effects of exercise: a critical review. Phys Sportsmed 1981;9:46–55
- Johnsgard KW. The Exercise Prescription for Anxiety and Depression. New York, NY: Plenum Publishing; 1989
- Morgan WP. Affective beneficience of vigorous physical activity. Med Sci Sports Exerc 1985;17:94–100
- Dishman RK. The norepinephrine hypothesis. In: Morgan WP, ed. Physical Activity and Mental Health. Washington, DC: Taylor & Francis; 1997
- Ebert MH, Post RM, Goodwin FK. Effect of physical activity on urinary MHPG excretion in depressed patients [abstract]. Lancet 1972;2:766
- Post RM, Kotin J, Goodwin FK, et al. Psychomotor activity and cerebrospinal fluid amine metabolites in affective illness. Am J Psychiatry 1973; 130:67–72
- 42. Tang SW, Stancer HC, Takahashi S, et al. Controlled exercise elevates plasma but not urinary MHPG and VMA. Psychiatry Res 1981;4:13–20
- Leith LM. Foundations of Exercise and Mental Health. Morgantown, WVa: Fitness Information Technology; 1994
- Bandura A. Self-Efficacy: The Exercise of Control. New York, NY: WH Freeman & Company; 1997
- Gleser J, Mendelberg H. Exercise and sport in mental health: a review of the literature. Isr J Psychiatry Relat Sci 1990;27:99–112
- Martinsen EW. Benefits of exercise for the treatment of depression. Sports Med 1990;9:380–389
- Morgan WP. Exercise and mental health. In: Dishman RK, ed. Exercise Adherence: Its Impact on Public Health. Champaign, Ill: Human Kinetics; 1988
- Raglin JS, Morgan WP. Influence of exercise and quiet rest on state anxiety and blood pressure. Med Sci Sports Exerc 1985;19:456–463
- Bortz WM, Angwin P, Mefford IN. Catecholamines, dopamine, and endorphin levels during extreme exercise. N Engl J Med 1981;305: 466–467
- Carr DB, Bullen BA, Skrinar G, et al. Physical conditioning facilitates the exercise-induced secretion of beta-endorphin and beta-lipotropin in women. N Engl J Med 1981;305:597–617
- Farrell PA, Gates WK, Maksud MG. Increases in plasma beta-endorphin/ beta-lipotropin immunoreactivity after treadmill running in humans. J Appl Physiol 1982;52:1245–1249
- Markoff RA, Ryan P, Young T. Endorphins and mood changes in long-distance running. Med Sci Sports Exerc 1982;14:11–15
- McMurray RG, Berry MJ, Hardy CJ. Physiologic and psychologic responses to a low dose of naloxone administered during prolonged running. Ann Sports Med 1988;4:21–25
- 54. Ransford CP. A role for amines in the antidepressant effect of exercise: a review. Med Sci Sports Exerc 1982;14:1–10
- Dunn AL, Reigle TG, Youngstedt SD, et al. Brain norepinephrine and metabolites after treadmill training and wheel running in rats. Med Sci Sports Exerc 1996;28:204–209
- Jacobs BL. Serotonin, motor activity and depression-related disorders. Am Sci 1994;82:456–463
- Martinsen EW. The role of aerobic exercise in the treatment of depression. Stress Med 1987;3:93–100
- Morrow J, Nolen-Hoeksema S. Effects of responses to depression on the remediation of depressive affect. J Pers Soc Psychol 1990;58:519–527
- Nolen-Hoeksema S, Morrow J. A prospective study of depression and posttraumatic stress symptoms after a natural disaster: the 1989 Loma Prieta earthquake. J Pers Soc Psychol 1991;61:115–121
- Klein MH, Greist JH, Gurman AS, et al. A comparative outcome study of group psychotherapy vs. exercise treatments for depression. Int J Ment Health 1985;13:148–177
- Brown SW, Welsh MC, Labbe EE, et al. Aerobic exercise in the psychological treatment of adolescents. Percept Mot Skills 1992;74:555–560
- 62. Craft LL. Exercise and clinical depression: examining two psychological mechanisms. Psychol Sport Exerc. In press
- 63. American College of Sports Medicine. The recommended quantity and quality of exercise for developing and maintaining cardiorespiratory and muscular fitness, and flexibility in healthy adults: position stand. Med Sci Sports Exerc 1998;30:975–991

- Berger BG, Owen DR. Relation of low and moderate intensity exercise with acute mood change in college joggers. Percept Mot Skills 1998;87: 611–621
- Hardy CJ, Rejeski WJ. Not what, but how one feels: the measurement of affect during exercise. J Sport Exerc Psychol 1989;11:304–317
- 66. McGowan RW, Talton BJ, Thompson M. Changes in scores on the Profile of Mood States following a single bout of physical activity: heart rate and changes in affect. Percept Mot Skills 1996;83:859–866
- Moses J, Steptoe A, Mathews A, et al. The effects of exercise training on mental well-being in the normal population: a controlled trial. J Psychosom Res 1989;33:47–61
- Parfitt G, Eston R, Connolly D. Psychological affect at different ratings of perceived exertion in high-and low-active women: a study using a production protocol. Percept Mot Skills 1996;82:1035–1042
- Steptoe A, Cox S. Acute effects of aerobic exercise on mood. Health Psychol 1988;7:329–340
- Sallis JF, Haskell WL, Fortmann SP. Predictors of adoption and maintenance of physical activity in a community sample. Prev Med 1986;15: 331–334
- Driver HS, Taylor SR. Exercise and sleep. Sleep Med Rev 2000;4: 387–402
- 72. Youngstedt SD, O'Connor PJ, Dishman RK. The effects of acute exercise on sleep: a quantitative synthesis. Sleep 1997;20:203–214

- Buckworth J, Wallace LS. Application of the transtheoretical model to physically active adults. J Sports Med Phys Fitness 2002;42:360–367
- Duncan KA, Pozehl B. Staying on course: the effects of an adherence facilitation intervention on home exercise participation. Prog Cardiovasc Nurs 2002;17:59–65
- 75. King AL, Taylor CB, Haskell WL, et al. Strategies for increasing early adherence to and long-term maintenance of home-based exercise training in healthy middle-aged men and women. Am J Cardiol 1988;61:628–632
- 76. Writing Group for the Activity Counseling Trial Research Group. Effects of physical activity counseling in primary care: the Activity Counseling Trial: a randomized controlled trial. JAMA 2001;286:677–687
- Albright CL, Cohen S, Gibbons L, et al. Incorporating physical activity advice into primary care: physician delivered advice within the activity counseling trial. Am J Prev Med 2000;18:225–234
- Calfas KJ, Long BJ, Sallis JR, et al. A controlled trial of physician counseling to promote the adoption of physical activity. Prev Med 1996;25: 225–233
- Green BB, McAfee T, Hindmarsh M, et al. Effectiveness of telephone support in increasing physical activity levels in primary care patients. Am J Prev Med 2002;22:177–183
- Pinto BM, Friedman R, Marcus BH, et al. Effects of a computer-based telephone counseling system on physical activity. Am J Prev Med 2002;23:113–120

For the CME Posttest for this article, see pages 112–113.



Instructions and Posttest

Participants may receive up to 1 Category 1 credit toward the American Medical Association Physician's Recognition Award by reading the CME article and correctly answering at least 70% of the questions in the Posttest that follows:

Go to **www.psychiatrist.com/cmehome** to take this Posttest online and earn credit immediately.

Or

- 1. Read each question carefully and circle the answer on the Registration Form.
- 2. Type or print the registration information in the spaces provided and complete the evaluation.
- 3. Send the Registration Form to the address or fax number listed on the Registration Form.

All replies and results are confidential. Answer sheets, once graded, will not be returned. Unanswered questions will be considered incorrect and so scored. The CME Institute of Physicians Postgraduate Press, Inc. will keep only a record of participation, which indicates the completion of the activity and the designated number of Category 1 credits that have been awarded. Correct answers to the Posttest will be made available to the participants of this activity upon request after the submission deadline.

Accreditation Statement

Physicians Postgraduate Press, Inc. is accredited by the Accreditation Council for Continuing Medical Education to provide continuing medical education for physicians.

- 1. According to meta-analytic findings, the only exercise program characteristic to significantly moderate the effect was _____.
 - a. Frequency of exercise
 - b. Duration of the exercise bout
 - c. Intensity of the exercise bout
 - d. Length of the exercise program
- Results from meta-analytic studies suggest that an increase in success rate due to treatment (i.e., exercise) can be as high as _____% for clinically depressed patients.
 a. 74
 - b. 25
 - c. 10
 - d. 35
- 3. Which of the following proposed mechanisms for the exercise-depression relationship suggests that exercise leads to cognitive changes in perspective and confidence? a. Endorphin hypothesis
 - b. Monoamine hypothesis
 - c. Thermogenic hypothesis
 - d. Self-efficacy hypothesis
- An initial exercise prescription of _____ min per day, _____ times per week, at _____ intensity is recommended for depressed patients.
 - a. 45, 5, moderate
 - b. 20, 3, moderate
 - c. 45, 3, vigorous
 - d. 20, 5, vigorous
- 5. Which of the following techniques has been associated with greater exercise adherence?
 - a. Have patients self-monitor their progress with exercise logs or pedometer counts
 - b. Have physicians track patients' exercise by monitoring changes in weight
 - c. Provide patients with information regarding negative health consequences of inactivity
 - d. None of the above have been associated with exercise adherence

Answer to Pretest: 1. b



Registration and Evaluation

Circle the one correct answer for each question. 4. 1. а b с d а b с d 5. 2. b с d а b а с d 3. а b d С Print or type Name Social Security number _____ – _____ (for CME credit recording purposes) Degree _____Specialty _____ Affiliation Address City, State, Zip _____ Phone ()_____ Fax () E-mail Hospital Private Practice Resident 🖵 Intern

Deadline for submission

For a credit certificate to be issued, please complete this Registration Form no later than June 30, 2006. Online submissions will receive credit certificates immediately. Faxed or mailed submissions will receive credit certificates within 6 to 8 weeks.

Keep a copy for your files

Retain a copy of your answers and compare them with the correct answers, which will be published after the submission deadline.

Payment

If you complete the test online, no payment is necessary. A \$10 payment must accompany this form. You may pay by check, money order, or credit card (Visa or MasterCard). Make check or money order payable to Physicians Postgraduate Press, Inc. If paying by credit card, please provide the information below.

Check one: 🗅 Visa 🖨 MasterCard	
Card number	10. How much ti CME activity
Expiration date	11. Do you have
Your signature	\Box Yes \Box No

Please evaluate the effectiveness of this CME activity by answering the following questions.

- 1. Was the educational content relevant to the stated educational objectives? □ Yes □ No
- 2. Did this activity provide information that is useful in your clinical practice? □ Yes □ No
- 3. Was the format of this activity appropriate for the content being presented? □ Yes □ No
- 4. Did the method of presentation hold your interest and make the material easy to understand?
 □ Yes □ No
- 5. Achievement of educational objective:
 - A. Enabled me to counsel patients on the adoption of exercise as an adjunct treatment for depression. □ Yes □ No
- 6. Did this CME activity provide a balanced, scientifically rigorous presentation of therapeutic options related to the topic, without commercial bias? □ Yes □ No
- 7. Does the information you received from this CME activity confirm the way you presently manage your patients? □ Yes □ No
- 8. Does the information you received from this CME activity change the way you will manage your patients in the future? □ Yes □ No
- 9. Please offer comments and/or suggested topics for future CME activities.
- 10. How much time did you spend completing this CME activity?
- 11. Do you have convenient access to the Internet?
 □ Yes □ No

TEAR OUT AND SEND THIS PAGE, ALONG WITH YOUR PAYMENT, TO:

CME INSTITUTE • PHYSICIANS POSTGRADUATE PRESS, INC. • P.O. BOX 752870 • MEMPHIS, TN 38175-2870 IF YOU ARE PAYING BY CREDIT CARD, YOU MAY FAX THIS PAGE TO: CME INSTITUTE AT 901-751-3444 QUESTIONS? CALL 1-800-489-1001 EXT. 8 • WWW.PSYCHIATRIST.COM