LETTER

Exercise is medicine for depression: even when the "pill" is small

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Dear editor

In a recent perspectives article, Gerber et al argue persuasively that more systematic efforts are needed to utilize the considerable treatment potential of exercise in psychiatric care.¹ Specifically, the authors highlight the inherent difficulties that clinicians and researchers face when encouraging patients to initiate and maintain a regular physical activity program. These obstacles relate to dysfunctional cognitive– emotional processes which, the authors argue, can interfere with patients' capacity to self-regulate health-related behaviors such as exercise. They further suggest that behavioral skill training should be used to support patients with major depressive disorders (MDDs), and that the assessment of physical activity and cardiorespiratory fitness should be routine in psychiatric practice. Finally, the authors comment on the optimal exercise dose, and draw attention to the paucity of studies exploring the long-term effects of exercise on depression. We commend the authors for their timely recommendations and concur with their main suggestions. In this brief letter, we wish to complement Gerber's excellent article with three key points.

First, we wish to reemphasize the importance of interventions that aim to improve the physical health of those with depression. While suicide accounts for some of the premature deaths among people with MDDs, it is now established that high levels of cardiovascular and metabolic disease in this population^{2,3} are leading contributors to the premature mortality gap of 10–15 years in people with depression compared to the general population.⁴ It has also been shown that individuals who maintain their cardiorespiratory fitness across the lifespan have a reduced risk of experiencing a depressive episode,⁵ in addition to other detrimental health outcomes, such as cardiovascular disease and diabetes. Reducing these cardio-metabolic health problems is unlikely to be achieved as efficiently through antidepressant use or other psychological interventions alone.

Second, we agree that supported exercise interventions involving trained health professionals, such as exercise physiologists, psychologists and nurses, are necessary to optimize treatment outcomes. Recent meta-analyses have demonstrated that cardiorespiratory fitness gains from exercise are higher, and dropout levels lower, when recognized exercise professionals are used.^{6,7} Indeed a review of the program variables in exercise for depression studies concluded that individually tailoring exercise to suit the client's preference, access to facilities and ensuring supervision are ways to enhance treatment outcomes.⁸ In practical terms, this can be achieved through weekly face-to-face contact with a qualified health professional where goal setting is discussed and exercise benefits and barriers reviewed, or through "remote" contact

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using email and text messaging to encourage adherence to exercise regimes. While plausible, Gerber's suggestion that "major therapeutic efforts are required to decrease the unfavorable cognitive–emotional processes experienced by patients with MDDs" (p 1979) has yet to be empirically tested in the context of exercise training. Further research is needed to determine what constitutes optimal supervision of this patient group.

Finally, we wish to comment on the "dose" of exercise needed to obtain clinically meaningful treatment effects. Gerber et al cite a previous efficacy study performed in a laboratory setting indicating that exercise needs to be performed at "public health doses" to achieve a treatment effect.9 While this groundbreaking and well-executed study remains relevant today, findings from effectiveness trials can also inform this discussion. For example, in a recent community-based trial, Hallgren et al randomized 946 outpatients with mild-to-moderate depression to one of three 12-week interventions: supervised physical exercise, internet-based cognitive behavioral therapy (CBT) and usual care by a physician, consisting of CBT-focused therapy and antidepressant treatment.¹⁰ Participants in the exercise condition were further randomized to one of three exercise intensities: light exercise consisting of yoga and stretching (without a mindfulness component), moderate aerobic exercise or vigorous aerobic exercise. All sessions were calibrated and conducted in established fitness facilities with qualified instructors. Although participants were encouraged to complete three exercise sessions per week (36 in total), most only attended about 1 exercise class per week. At posttreatment, patients randomized to the exercise intervention reported significantly lower depression severity compared to usual care. Of importance, the positive effects of exercise on depression severity were maintained 12 months after the baseline assessment, despite suboptimal adherence.¹¹ Moreover, the effects of exercise intensity were equivalent across groups, suggesting that all three intensities were equally effective in the treatment of mild-to-moderate depression.12

In summary, we concur with and thank Gerber et al for their important recommendations. We also wish to highlight promising new results from community-based trials with longterm follow-up assessments indicating that exercise is indeed medicine for depression, even when the "pill" is small.

Disclosure

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Dear editor

In their letter, Hallgren, Vancampfort and Stubbs supported our claims¹ that more systematic efforts are required to increase physical activity and cardiorespiratory fitness in psychiatric settings, and to better utilize the potential of exercise as a treatment in psychiatric care. They also shared our view that behavior skill training may be a promising avenue to increase exercise adherence. In this respect, Mota-Pereira et al² showed that regular and moderately intensive walking improved symptoms of depression in treatment-resistant patients with major depressive disorders. Specifically, they employed both supervised and non-supervised techniques. The latter consisted of reminders to exercise by placing sports shoes by the front door, leaving a note on the main door or using cell phone reminders with a specific ringtone. Further, "social enhancers" were employed; patients were encouraged by and involved family members, in order to both remind them to be physically active and to exercise together with them. We hold that such pragmatic tricks are more efficient, compared to simply advising patients to increase their cardiorespiratory fitness levels. Further, we claim that "cardiorespiratory fitness" seems to be a rather academic, intellectual and "bulky" concept, which may be difficult to translate to patients in a comprehensible form (How does cardiorespiratory fitness feel, and what should I do to improve it?). Accordingly, we claim that the "next generation" of exercise interventions targeting patients with major depressive disorders should take into account which environmental, social and above all cognitive-emotional processes might boost and stabilize the social and cognitive determinants of exercise regulation. As a result of this, questions arise as to why it seems so difficult, even for people without psychiatric diagnoses, to maintain regular exercise regimes, resulting in the fact that physical inactivity is among the main causes of both somatic and psychiatric diseases worldwide.^{3–5}

We thank Hallgren, Vancampfort and Stubbs for their appreciatory comments and for sharing their own ideas. Hallgren et al complemented our perspectives with several important points, which we would like to comment on. In particular, they emphasized that the effectiveness of behavioral skill training has not yet been tested in patients with psychiatric disorders and that even small doses of physical activity can lead to lasting reductions in depressive symptoms among patients with mild-to-moderate depression. Here, again, we refer to the seminal work of Mota-Pereira et al.² Moreover, Chalder et al⁶ recently showed, in a study with 361 participants who had consulted their general practitioner with symptoms of depression, that a protocol of three face-to-face sessions and 10 telephone calls with a trained physical activity facilitator (over an 8-month period) was associated with increased physical activity levels at the 4-month follow-up (adjusted odds ratio: 2.27, P<0.01). Nevertheless, given that behavior skill training might be a special challenge in psychiatric care due to unfavorable cognitive-emotional processes experienced by these patients (eg, hopelessness, pessimism, lack of motivation), we believe that more well-controlled intervention studies are needed to examine whether a generalization to psychiatric populations is possible or whether alternative approaches must be taken into consideration.

Hallgren et al's second claim (that even small doses of physical activity are able to decrease depressive symptoms) seems justified. Based on a study with 946 outpatients who were randomly assigned to an internet-based cognitive therapy, usual care by the physician or light, moderate or vigorous aerobic exercise, they concluded that after the end of the intervention phase, patients of all exercise groups reported lower levels of depressive symptoms compared to usual care, an effect which was maintained until 12 months after baseline assessment.⁷⁻⁹ In summary, these findings suggest that far less exercise is needed to decrease depressive symptoms than typically suggested in internationally accepted physical activity recommendations.^{10,11} Further, this pattern of results mirrors the findings of Mota-Pereira et al² among treatment-resistant patients with major depressive disorders: while after 12 weeks, none of the 10 patients in the control condition showed any improvement (but also no aggravation), of the 19 patients in the walking group, 10 (52.6%) remained in a state of depression, 4 patients (21%) were responders and 5 (26.3%) were fully remitted. Or simply put, in the walking group, improvements were

observed in half of the patients with treatment-resistant major depression.

With regard to the question of how the "next generation" of research and interventions might look in this field of study, we would like to emphasize seven points:

First, we question the extent to which most physical activity recommendations are appropriate for patients with psychiatric disorders, as physical activity recommendations have generally been developed with a focus on the prevention of cardiovascular diseases.^{12,13} In this regard, in a recent study with patients suffering from stress-related exhaustion disorder,¹⁴ we observed the following results: in originally inactive patients, compliance with physical activity recommendations 6 months after the end of multimodal treatment was associated with continued decreases in depression and burnout symptoms. Next, not surprisingly, continued physical inactivity was linked with a new rise in symptom severity. However, similar to Hallgren et al^{7,8} and Helgadóttir et al,⁹ we found that patients who only exercised 1-2 times per week (and thus, did not meet recommended physical activity standards) achieved comparable benefits to their counterparts with more frequent exercise involvement.¹⁴ In summary, these findings suggest that psychological symptoms can be prevented or improved with small amounts of exercise, below the levels recommended to improve cardiovascular health.^{15–17} Some of our studies even suggest that the way in which people think about their physical activity levels is more closely related to their psychological functioning than their self-reported exercise.^{18,19} However, since it is reasonable to assume that such minor doses of exercise are not sufficient to improve cardiorespiratory fitness, and thus have only a limited potential to enhance physical health and to reduce the premature mortality rate in people with psychiatric disorders, meeting current physical activity recommendations still seems to be a meaningful target for psychiatric patients.

Second, at a psychological level and from a self-efficacy perspective, we question the cognitive-emotional importance of the concept of "cardiorespiratory fitness" to motivate psychiatric patients to adopt more physically active lifestyles. We claim that this concept is too abstract, and that even for people without psychiatric issues, "cardiorespiratory fitness" is difficult to capture; accordingly, how should people aim for targets of which they have no (clear) mental representation? Poorly defined aims lack the capacity to improve goaloriented and behavior-steering cognitions.^{20–22} More specifically, we claim that any kind of physical activity and fitness target should be visualized so that patients with psychiatric issues can literally see their improvements.²³ To this end, graphs, figures and appropriate technological tools are helpful, albeit not sufficient. We hold that encouragement through well-trained professionals is of key importance because patients with psychiatric disorders tend to bias and downplay improvements, or attribute any kind of improvement to external sources.²⁴ Thus, instructors should emphasize that improvements with regard to physical activity and fitness are exclusively related to patients' efforts, hereby improving patients' self-efficacy. Moreover, we posit that doses of exercise should be increased gradually in order to avoid feelings of failure and helplessness, which might be counterproductive in the long run.²⁵ Such an approach is also mirrored in the behavioral analysis as proposed by Kanfer and Saslow.²⁶ Specifically, the authors highlight the importance of improving patients' self-observation and shifting self-evaluation from an unfavorable to a favorable evaluation.

Third, we strongly believe that the promotion of physically active lifestyles in people with psychiatric disorders needs professional stimulation and supervision, along with the training of practical behavioral techniques that facilitate exercise adherence, as impressively shown by Mota-Pereira et al in patients with treatment-resistant disorders,² by Chalder et al in people diagnosed with an episode of depression⁶ and by many other researchers with other patient groups.^{27–35}

Fourth, looking back, many previous studies have focused on short-term effects of exercise interventions, placing a particular focus on psychological symptoms, cardiorespiratory fitness and/or physiological mechanisms as outcome variables. Moreover, in an attempt to increase internal consistency and to draw conclusions about cause and effect, many of these studies have been carried out in strongly controlled settings. While these studies have laid important groundwork by raising our awareness that (a) in the short run, exercise activities and usual care or psychopharmacological treatment are similarly effective, and (b) exercise may be used as a valuable add-on to standard treatment, few efforts have been made to examine long-term effects. Nevertheless, many of these studies were atheoretical and have failed to integrate current knowledge from established behavior change models. As a consequence, few studies have tested whether the interventions had a positive impact on social, cognitive and affective factors involved in the regulation of physical activity and exercise behavior.³⁶ Given this background, we reemphasize our claim that more systematic and theory-based efforts are needed to enable (and assess) long-term behavior change.37,38

Fifth, while we believe that motivational and volitional interventions are well suited to addressing explicit

motivational processes such as self-efficacy beliefs, outcome expectancies, behavioral intentions, implementation intention or coping skills, sport and exercise scientists have recently highlighted the relevance of implicit attitudes in the regulation of exercise behavior. For instance, Conroy et al³⁹ showed in a sample of healthy participants that implicit attitudes contributed to the prediction of physical activity beyond well-established predictors of intentional physical activity. Conroy et al therefore concluded that "physical activity motivation involves both explicit and implicit processes, and physical activity promotion efforts may be enhanced by attending to relevant implicit motivation processes" (p 112). Similar findings were reported by Bluemke et al⁴⁰ and Chevance et al.⁴¹ Importantly, Antoniewicz and Brand⁴² have shown that among undergraduate students, implicit attitudes toward exercise can be improved through computer-based evaluative conditioning and that this procedure can move people toward developing an enjoyment of exercise. Given this background, future research could address the question of whether psychiatric patients and healthy controls differ with regard to their automatic evaluations of exercise. If this were the case, fostering simultaneously explicit and implicit motivational processes to promote increased lifestyle physical activity would seem a worthwhile endeavor in psychiatric care.

Sixth, Mothes et al⁴³ showed that in healthy adult exercisers who believed that regular exercising was healthy, exercise had a stronger impact on both subjective and neurophysiological health outcomes than in participants who were less convinced of the health-enhancing potential of regular exercise. In patients with psychiatric disorders, it might be especially challenging to overcome negative automatic evaluations of exercise. We therefore emphasize that providers of physical activity and exercise programs should be aware that the affective responses during and after exercise will have a significant impact on whether participants will adhere to regular physical activity.44,45 Hence, in line with the work of Ekkekakis et al,^{46–48} we suggest that efforts are needed to make exercise for patients with psychiatric disorders as pleasant and enjoyable as possible, for example, by choosing the right activities and intensities or creating an autonomy-supportive climate.49,50

Seventh, we claim that more studies in naturalistic settings or based on patient registers are needed to find out whether the existing positive effects of exercise on psychological functioning (based on randomized controlled trials) can be replicated if physical activity and exercise programs are integrated in existing clinical structures. Finally, we claim that more information is required with regard to the costeffectiveness of exercise and physical activity (counseling) programs in psychiatric care in order to compare exercise therapy with more traditional treatments.

Again, we thank Hallgren et al for their important comments, and hope that our response will stimulate new and fruitful debates regarding the question of how "next generation" research and intervention programs may look in this field of research.

Disclosure

The authors report no conflicts of interest in this communication.

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