

Physical exercise and migraine: for or against?

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Introduction

Migraine can now be considered a major public healthcare issues, since it is one of the leading causes of disability worldwide, causing a considerable number of clinical, social and economic issues (1). Recent statistics attest that the prevalence of this condition may be as high as 25% in adults, with approximately 4% of the general population suffering from a chronic form of disease. Albeit the precise pathogenesis of migraine still remains substantially unknown (2), many effective therapeutic strategies have been proposed, mainly entailing the administration (alone or in combination) of triptans, ergots, acetaminophen and nonsteroidal anti-inflammatory drugs (NSAIDs). Besides, there is still uncertainty of the real effectiveness in many migraine patients of some possible prophylactic or preventive measures encompassing drugs, lifestyle changes and dietary habits (1).

The relationship between physical exercise and migraine remains rather controversial. On one hand, worsening of migraine pain has often been perceived by patients during or after regular physical exercise (3,4). However, in a critical review of scientific literature published nearly 10 years ago, Busch and Gaul concluded that although the data available at that time was mostly insufficient for establishing whether or not exercise may be an effective therapeutic or preventive measure for improving migraine symptoms and quality of life of affected patients, an active lifestyle is often recommended by many migraine experts (5). Ten years afterward, many and more recent epidemiologic and intervention studies were published, which should actually provide an updated and more straightforward

contribution to dispel the doubts about the intriguing relationship between physical exercise and migraine. Therefore, the aim of this article is to provide a narrative review of recent scientific articles (i.e., published from the year 2008 to present time) which have explored the potential association between physical exercise and migraine.

Epidemiologic studies

Varkey *et al.* carried out a large prospective study including 22,397 participants to investigate the potential relationship between physical exercise and migraine (6). After 11 years of follow-up, a light physical exercise (i.e., no sweating or being out of breath) was associated with a 35% enhanced risk of migraine (OR, 1.35; 95% CI, 1.21–1.51) compared to hard physical exercise (i.e., sweating or being out of breath).

Milde-Busch *et al.* administered a lifestyle questionnaire to 1,260 adolescents to identify potential correlates of migraine (7). These authors found that a low level of metabolic equivalent index of physical exercise was associated with a 4.2-fold higher risk of migraine compared with to a high level of metabolic equivalent index [odds ratio (OR), 4.2; 95% confidence interval (95% CI), 2.2–7.9].

Winter *et al.* reported the results of a large population study based on 7,417 individuals, in which the potential relationship between many lifestyle factors and migraine was investigated (8). No association was observed between the level of physical exercise and the risk of developing migraine attacks.

In a subsequent article, Winsvold *et al.* published the

results of the Nord-Trøndelag Health Study in Norway 1995–1997 (HUNT 2) study, in which 48,713 subjects aged 20 years or older were administered with a questionnaire on potential lifestyle factors associated with development of migraine (9). Compared to patients without migraine (63.6%), those with (68.7%) or without (68.4%) aura reported a significantly higher burden of physical inactivity.

Koppen *et al.* interviewed 103 migraine patients first presenting at a headache clinic to identify potential correlates of migraine (10). Overall, 39/103 patients (38%) reported a risk of developing migraine attacks during exercise, most of which during endurance activities (i.e., running). All of these patients also reported that migraine attacks tended to worsen by practicing physical exercise. Interestingly, neck pain and headaches were both more frequent in patients with exercise-provoked migraine than in those without.

Lebedeva *et al.* performed a large population study, including 3,124 subjects (43% women) who were interviewed with the purpose of identifying potential lifestyle factors associated with the risk of developing migraine (11). In multivariate logistic regression analysis, low physical exercise (i.e., <30 min of physical exercise per week) was associated with a 60% higher risk of developing migraine (OR, 1.6; 95% CI, 1.0–2.4). Notably, this association was found to be statistically significant in women (OR, 1.9; 95% CI, 1.1–3.2) but not in men (OR, 1.2; 95% CI, 0.6–2.6).

Krøll *et al.* performed a population study based on 148 patients followed at a tertiary specialized migraine center and 100 healthy matched controls, who were interviewed about their lifestyle factors (12). The risk of having a migraine was hence found to be 4.4-fold higher (OR, 4.4; 95% CI, 1.6–11.7) for subjects with a low level of physical exercise (<30 min of moderate-intensity physical exercise on most days) compared to those with a high level (≥ 1 hour per day or more of moderate-intensity physical exercise).

Similar data was reported in a subsequent study by Farris *et al.*, who administered a physical activity questionnaire to 132 migraine women aged between 18–50 (13). According to the results of the surveys, no significant variation of pain while performing physical activity was reported in the vast majority of participants (i.e., 62%), worsened pain was reported by 35% of participants, whilst symptoms improvement was only noticed by 3% of participants. Notably, physical activity led to pain worsening during all attacks in only 10% of participants.

Hagen *et al.* carried out a population-based historical cohort study (Nord-Trøndelag Health Study) to investigate the impact of different lifestyle aspects on the risk of developing migraine (14). The study population consisted of 15,276 participants without migraine at baseline, who were then evaluated over a period of 11 years. Interestingly, 1 to 3 hours of low-intensity physical exercise per week and 1 to 2 hours of hard physical exercise per week were respectively associated with a 22% [relative risk (RR), 0.78; 95% CI, 0.62–0.99] and 29% (RR, 0.71; 95% CI, 0.54–0.94) lower risk of developing migraine compared to inactivity. A trend towards lower risk of developing migraine was also observed for ≥ 3 hours of high-intensity physical exercise per week, but the association did not reach statistical significance (RR, 0.92; 95% CI, 0.67–1.27).

In another study, Luedtke *et al.* explored surface electromyography responses of trapezius muscle in 28 healthy subjects as well as in 31 patients with chronic migraine and 43 with episodic migraine (15). Increased bilateral trapezius muscle activity was recorded in all groups during physical exercise, but no statistically significant difference could be observed ($P < 0.05$). It could hence be concluded that trapezius muscle activity should not be considered an underlying trigger but rather should be interpreted as an accompanying symptom in migraineurs.

Intervention studies

Albeit published before 2008, the article of Narin *et al.* deserves to be mentioned, since it is one of most accurate studies exploring the effect of physical exercise on migraine (16). Briefly, 40 migraine women were followed by the neurology department and were equally randomized to an exercise group undertaking 1-hour aerobic exercise 3 times weekly and to a control group. After 8 weeks, the intervention group had a substantial improvement of all migraine features, thus displaying a reduction of pain intensity (–55% versus pre-exercise and –43% versus control group), pain duration (–67% versus pre-exercise and –58% versus control group), pain frequency (–51% versus pre-exercise and –49% versus control group) and pain disability index (–56% versus pre-exercise and –51% versus control group).

In a subsequent study, Dittrich *et al.* equally randomized 30 migraine women to an aerobic exercise group (1-hour of twice-weekly indoor aerobic exercise program for 6 weeks) and a control group (no change of previous

physical exercise level) (17). After the intervention period a substantial reduction of intense migraine pain (-80% versus pre-exercise and -75% versus control group) and frequency of migraine attacks (≥ 1 per week; -50% versus pre-exercise and -67% versus control group) could be recorded. Interestingly, nearly 90% of patients in the exercise group perceived that physical exercise was helpful to ameliorate active behavior.

Varkey *et al.* carried out an intervention study based on 26 migraine patients and entailing 40 min indoor cycling 3 times weekly for 12 weeks (18). At the end of treatment, a substantial improvement could be observed, as attested by a reduction of pain intensity (-18%), migraine attacks per month (-23%), days with migraine (-28%) and use of anti-migraine drugs (-24%). Notably, the perceived quality of life was also significantly improved after treatment ($+12\%$).

Pairo and collaborators performed a random semi-experimental study including 20 migraine women (19). The women were randomized into an exercise group (i.e., 40 min of moderate intensity aerobic exercise 3 times weekly) and a control group (no exercise). One month after the intervention, frequency, intensity and duration of migraine attacks were found to be substantially reduced.

Varkey and coworkers carried out another study based on a specific subset of 14 migraine patients, reporting that physical exercise often led to pain worsening during a migraine attack (20). All patients underwent two exhaustion cycle ergometer tests. Overall, a migraine attack developed after both tests in 3/14 patients (21%), after one test in 5/14 patients (36%), whilst 6/14 patients (43%) did not develop a migraine attack after either trial. Notably, a modest but significant association ($P=0.036$) was found between test-provoked migraine and baseline frequency of migraine attacks.

Hanssen *et al.* performed a controlled randomized trial involving episodic migraine patients, who were randomized into three groups, i.e., high-intensity interval training (HIT; 4 series of 1 min treadmill exercise at heart rate between 90–95% followed by 3 min treadmill exercise at target heart rate of 70%, twice weekly), moderate continuous exercise training (MCT; 45 min treadmill exercise at target heart rate of 70%, twice weekly) and no exercise (21). After a 12-week period, both types of exercise were associated with a reduced frequency of migraine attacks, although HIT seemed globally more efficient (-63%) than MCT (-26%).

Interesting findings emerged from a recent intervention

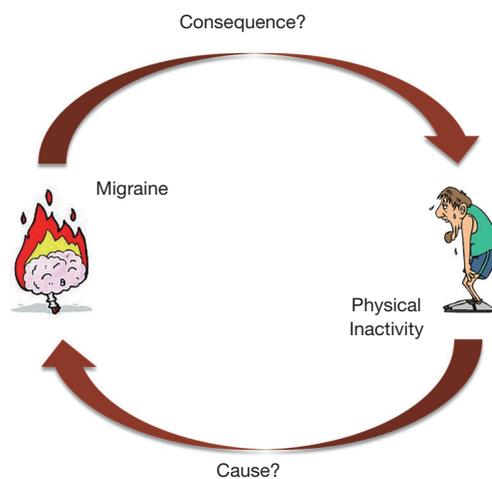
study, in which 52 migraine patients were equally randomized into an exercise group (i.e., 45 min of aerobic exercise 3 times weekly for 3 months) and a control group (i.e., usual daily activities, no aerobic physical exercise) (22). After the end of the study and during 6 months of follow-up, the aerobic exercise group displayed a substantial improvement of all the main migraine features, i.e., reduction of days with migraine (-23% versus both pre-exercise and control group), pain intensity (-16% versus pre-exercise and -8% versus control group) and pain duration (-28% versus pre-exercise, no difference with control group). Even more importantly, the quality of life of the exercise group was substantially improved during follow-up, as shown by a remarkable increase in the time spent on household chores ($+32\%$), paid work or study tasks ($+46\%$), social activities ($+31\%$), family life ($+33\%$), and physical exercise ($+17\%$).

Conclusions

The current available evidence garnered from the respectable number of recent epidemiologic investigations and intervention studies would lead us to formulate a number of evidence-based conclusions. Indeed, the results of most recent epidemiological studies are consistent to show an inverse association between physical exercise and migraine, but it is quite challenging to establish whether physical inactivity may be the cause or the consequence of migraine, since migraine is a well-established disabling condition that may actually limit social and physical capabilities (23). The evidence that emerged from the noticeable number of recent intervention studies seems more solid and reliable. Overall, a general reduction of frequency, severity and duration of migraine attacks has been observed after the exercise intervention periods, and this has also been frequently associated with a considerable improvement of the quality of life of the affected subjects (Table 1). These findings may be of substantial value in a clinical perspective. Severe migraine has a strong and unfavorable impact on the quality of life of the affected patients, since it is a disabling condition which not only impairs social life, but also unfavorably impacts physical functioning (24), often leading to compromised postural control performance (25). This evidence would hence pave the way to hypothesizing the existence of a vicious cycle, in which migraine becomes a major impediment for regular exercise performance and, in turn, a lower

Table 1 Intervention studies which have explored the putative benefits of physical exercise on migraine

Authors	Type of study	Intervention	Outcome
Narin <i>et al.</i> , 2003 (16)	Randomized, intervention study	1 hour aerobic exercise 3 times weekly, for 8 weeks	Reduction of pain intensity, pain duration, pain frequency and pain disability index
Dittrich <i>et al.</i> , 2008 (17)	Randomized, intervention study	1 hour aerobic exercise program twice weekly, for 6 weeks	Reduction of both intense migraine pain and high frequency of migraine attacks; improvement of active behavior
Varkey <i>et al.</i> , 2009 (18)	Intervention study	40 min indoor cycling 3 times weekly, for 12 weeks	Reduction of pain intensity and frequency of migraine attacks; lower use of anti-migraine drugs; improvement of perceived quality of life
Pauro <i>et al.</i> , 2016 (19)	Randomized, intervention study	40 min moderate intensity aerobic exercise 3 times weekly, for 1 month	Reduction of frequency, intensity and duration of migraine attacks
Varkey <i>et al.</i> , 2017 (20)	Intervention study	Double exhaustion cycle ergometer test	Development of migraine attack in 57% of patients with a history of exercise-provoked migraine
Hanssen <i>et al.</i> , 2017 (21)	Randomized, intervention study	4 series of 1 min treadmill exercise (heart rate 90–95%) followed by 3 min treadmill exercise (heart rate 70%) twice weekly; or 45 min treadmill exercise (heart rate 70%) twice weekly	Reduced frequency of migraine attacks
Krøll <i>et al.</i> , 2018 (22)	Randomized, intervention study	45 min of aerobic exercise 3 times per week, for 3 months	Reduction of pain intensity, pain duration and frequency of migraine attacks; improvement of both perceived quality of life and physically active lifestyle

**Figure 1** The “vicious cycle” between physical inactivity and migraine.

level of physical exercise would then contribute to worsen severity, frequency and duration of migraine attacks, thus further limiting the willingness of migraineurs to exercise (Figure 1). Therefore, regular engagement in moderate-intensity aerobic exercise (i.e., ≥ 40 min, 3 times per week) may be regarded as a reasonable preventive

measure in these subjects (Table 2). Many biological effects can be triggered by physical exercise, some of which may have a role in the pathogenesis of migraine. These will probably be mediated by chronic adaptation towards enhanced pain threshold and lower stress as commonly induced by regular physical exercise (26). This recommendation is indirectly supported by the evidence that even physiotherapy interventions (including manual therapy, soft-tissue technique and training) may be effective in reducing frequency and pain intensity of migraine (27). Whilst theoretically straightforward, this advice would not apply to patients with a personal history of exercise-provoked migraine (Table 2). This is due to the tangible risk that physical exercise may trigger or worsen a migraine attack in some patients, a process probably mediated by acute increase of the powerful vasodilation compound nitric oxide (NO) (16), which would then overcome the potential long-term benefits of physical exercise.

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None.

Table 2 Literature-based recommendations on physical activity and migraine

Regular moderate aerobic physical exercise (>40 min, 3 times per week) seems effective to reduce both severity and frequency of migraine attacks

Since exercising may sometimes worsen migraine, be engaged in physical exercise during a migraine attack must be established on an individual basis, according to the personal history of exercise-provoked migraine

High-intensity exercise should be avoided in patients with a history of exercise-provoked migraine

Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

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