TRIGGER FACTORS ASSOCIATED WITH MIGRAINE HEADACHE AMONG NORTHERN IRANIAN POPULATION

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ABSTRACT

Background. Migraine is the most common neurological disorder associated with a wide range of psychiatric co-morbidities. It is therefore regarded as a major global health problem due to its high prevalence. Here we investigated the factors associated with migraine among an Iranian population.

Methods. This investigation included a cross-sectional study consisting of 270 migraine patients. Initially, a questionnaire was designed based on demographic information and headache characteristics. SPSS analyzed the collected data, and chi-square and logistic regression were used to evaluate the clinical data.

Results. Our findings revealed that many migraine patients are experiencing multiple trigger factors, with stressful life events being the most common in both genders and migraine subtypes. The incidence of migraine was higher in women than in men. Among the trigger factors leading to the intensification of migraine attacks, stress, sleep changes, and bright lights were the most common and important causes of headache, followed by tiredness and weather change disturbance.

Conclusions. Considering the contradictory reports on the effect of trigger factors on migraine, further studies are needed to be performed. Due to the significance of the socio-economic impact of migraine on patient’s quality of life, strategies for managing these factors should be effectively implemented to improve the community’s health.

Keywords: headache, migraine disorders, symptoms, trigger factors

INTRODUCTION

Migraine is the most common primary headache disorder with episodic to chronic form demonstration (1). The headache disorder is characterized by unilateral, pulsating, recurrent attacks with moderate to severe pain intensity. Typically, symptoms linked include nausea, vomiting, photophobia, phonophobia, often leading to attacks that last 4-72 hours, and intensified by routine physical activity (2-4). The association between the prevalence of migraines and the socioeconomic status (SES) is unknown; however, based on clinical studies (5), it results in a significant disability with an enormous socioeconomic burden in addition to negatively affecting the person’s daily life, job performance and social activities, while depending on the level of severity, it can substantially impair quality of life (6,7). Migraine is the second headache disorder after tension-type headaches, estimated to affect approximately 15-18% of the general population (8,9). The prevalence is highly dependent on age and gender with more women (15-18%) being affected than men (6%) (10,11). According to epidemiological studies conducted in Asian countries, the prevalence of migraine has ranged from 8.4% to 12.7% (12). Some epidemio-
logical studies have shown that variables including, age, sex, premenstrual period, menstruation, sleep disorders, stress, skipping meals, level of education, income, climate, geographic region, nutrition, genetics, and environment affect migraine headaches to different degrees (13-16). Studies have shown that stress and other trigger factors can be closely associated with migraine attacks in susceptible patients, with anxiety affecting the severity and threshold of incidence and frequency of headaches (16,17). The links between food items and migraine headaches are complicated; however, the review of clinical scales implicate dietary triggers and the chemical constituent being involved in the mechanism of migraine attacks, affecting migraine’s pathophysiology at one or more phases of the migraine. Several different known migrainous triggers could influence the cerebral cortex, trigeminal nerve, brainstem trigeminal nuclei, thalamus, and brainstem or limbic pathways (17,18).

Before puberty, the prevalence of migraine is more frequently in boys than in girls, but after puberty, incidence and prevalence of headaches increase more quickly in girls than in boys, thence the prevalence increases in both genders by almost the age of 40 years, and then decreases (5,19). In many women with migraines, the menstrual cycle and stress are the main factors for migraine headaches. Studies have suggested that hormonal fluctuations, especially those of estrogen-related, are with migraines (20). Several studies on the factors affecting the exacerbation or prevalence of migraine attacks have been reported, with many factors not yet fully understood. Despite advances in the prevention and drug treatment of migraine, recognition of trigger factors, while remaining obscure, would provide an essential element for a successful clinical approach for headache patients, since avoiding such triggers reduces the severity and frequency of attacks are expected (16,21). According to a study in 2017 in Iran, the most common factors exacerbating migraine headaches included stress, bright lights, loud noise, fatigue, and overheating (22).

Epidemiological studies on the effect of headaches can help improve patient care and assess disease and correct classification. Due to different geographical distribution and different socio-cultural characteristics, the importance of epidemiological studies becomes apparent. This epidemiological study’s main purpose was to determine the associated factors of migraine headache, symptoms, and risk factors for disease progression of migraine in a northern Iranian population in a more comprehensive manner, with the hope of evaluating the existing conditions for future therapeutic interventions.

METHODS

Subjects

This study is of a cross-sectional type consisting of 270 migraine patients. From the patients referred to the Ayatollah Rouhani Hospital clinic in Babol-Iran, we received written informed consent to perform this study between October 2018 and December 2019. The diagnosis of migraine was according to the International Classification of Headache Society. Study inclusion criteria included patients referred to Ayatollah Rouhani Hospital, over the age of 16 and below 70-years-old, desire to participate in the study, lack of concomitant underlying diseases, and the exclusion criterion was an inadequate response to all the questionnaire objects. Also, we confirmed in interviews that migraine patients had no experience of medication overuse. For less than 12 months, patients with a headache disorder, with a primary or secondary headache, especially headache due to medication overuse, and severe psychiatric disease were excluded. The Ethics Committee approved the study protocol of the Kharazmi University, Tehran, Iran.

Study questionnaire

The method of data collection was the questionnaire. Questionnaires prepared for this study contained demographic information (age, gender, family history), headache characteristics (duration of symptoms, migraine whit aura, and migraine without aura), characteristics of accompanying symptoms (nausea, vomiting, photophobia, phonophobia), dietary factors (such as chocolate, nuts, tea/coffee, citrus fruits, fatty foods, spicy foods, meat), exertional activities, and potential risk factors for migraine attacks. The questionnaires and interviews were conducted face to face by neurologists and trained investigators.
Statistical analysis

The statistical analyses were conducted using SPSS software. Chi-squared test was used to compare the distributions of categorical variables between groups, and logistic regression was used for the evaluation of trigger factors. Odds ratios (OR) adjusted with 95% confidence intervals (CI) were calculated for trigger factors. P values < 0.05 were considered as statistically significant.

RESULTS

The migraine incidence was higher in females (92.2%) than in males (7.8%). The study population consisted of 270 patients, including 170 migraine without aura (158 females, 12 males), and 100 migraine with aura (91 females, nine males). A majority of the subjects were aged between 30 and 39 years (32.2%). Migraine was associated with a family history of migraine (60%). The frequency of headache was less than one attack per month in 13.7% of the participants, once a month is 20.4%, and more than one attack per month in 64.8% of the participants (Table 1).

TABLE 1. Demographic characteristics of study subjects showing prevalence rates of migraine

| Migraine prevalence rate | Number (%) |
|--------------------------|------------|
| Gender                   |            |
| Female                   | 249(92.2%) |
| Male                     | 21(7.8%)   |
| Age group (in years)     |            |
| <30                      | 63(23.3%)  |
| 30-39                    | 87(32.2%)  |
| 40-49                    | 75(27.8%)  |
| 50-59                    | 35(13.0%)  |
| 60-69                    | 10(3.7%)   |
| Family history           |            |
| Yes                      | 163(60%)   |
| No                       | 107(39.6%) |
| Frequency of headache    |            |
| Less than one attack per month | 37(13.7%) |
| Once a month             | 55(20.4%)  |
| More than one attack per month | 175(64.4%) |
| Headache quality         |            |
| Throbbing                | 212(78.5%) |
| Pressing                 | 58(21.5%)  |
| Aura                     |            |
| Yes                      | 100(37%)   |
| No                       | 170(63%)   |

Pain characteristics included 78.5% of migraineurs with headache reported as throbbing and 21.5% as pressing. Demographic characteristics of study subjects are shown in table 1. Among subjects suffering from migraine attacks, 46.6% of patients experienced photophobia symptoms during premonitory migraine phase. Among migraineurs, 37% experienced the aura phase of headache. Commonly associated symptoms with a headache have shown in figure 1. None of those mentioned above parameters demonstrated a statistically significant difference between migraine without aura and migraine with aura (Table 2). 64.1% of migraine patients experienced attacks with visual phenomena, while 36% experienced attacks with sensory aura. The most common associated symptoms during headache attacks were phonophobia (83.3%), photophobia (84.1%) and nausea (73.0%) (data have not shown).

TABLE 2. Demographic data and clinical characteristics in two study groups

| Headache type                      | Migraine without aura (N = 170) | Migraine with aura (N = 100) | p value |
|------------------------------------|---------------------------------|-----------------------------|---------|
| Migraine prevalence rate           |                                 |                             |         |
| Gender                             |                                 |                             |         |
| Female                             | 158(92.2)                       | 91(91.0)                    | 0.56    |
| Male                               | 12(7.1)                         | 9(9.0)                      |         |
| Age group (in years)               |                                 |                             |         |
| <30                                | 39(22.1)                        | 24(24.0)                    | 0.75    |
| 30-39                              | 51(30.0)                        | 36(36.0)                    |         |
| 40-49                              | 48(28.2)                        | 27(27.0)                    |         |
| 50-59                              | 25(14.7)                        | 10(10.0)                    |         |
| 60-69                              | 7(4.1)                          | 3(3.0)                      |         |
| Pain location                      |                                 |                             |         |
| On one side of head                | 68(40.1)                        | 49(49.0)                    | 0.57    |
| Whole head                         | 64(37.6)                        | 35(35.0)                    |         |
| Back                               | 4(2.4)                          | 2(2.0)                      |         |
| Front                              | 20(11.8)                        | 7(7.0)                      |         |
| Top                                | 14(8.2)                         | 7(7.0)                      |         |
| Family history                     |                                 |                             |         |
| Yes                                | 105(61.8)                       | 58(58.0)                    | 0.54    |
| No                                 | 65(38.2)                        | 42(42.0)                    |         |
| Headache quality                   |                                 |                             |         |
| Throbbing                          | 133(78.8)                       | 79(79.0)                    | 0.88    |
| Pressing                           | 37(21.8)                        | 21(21.0)                    |         |

According to the individuals’ experience, certain food types such as onion (42.6%), Sour food (39.3%), pepper (31.9%) and cheese (15.9%) intensified migraine headaches. Factors leading to the onset of migraine headaches included cigarette smoke (54%), stress (78.0%), sleep changes (69.6%), bright lights (57.8%), tiredness (55.6%) and weather changes (48.5%) (data not shown). The results on trigger factors and differences between gender types and subgroups of migraine are listed in tables 3 and 4. The most frequent trigger
factors among female patients with migraine included stress (79.9%), sleep changes (70%) and excitement (67.1), while those of male patients with migraine were stress (66%), fatigue (66%) and sleep changes (61%). A comparison of common triggers factors of headache with gender shown in Figure 2. Considering the P-values obtained from the logistic regression test; however, no significant correlation was detected between the trigger factors and intensification of migraine headache (Table 5).

**TABLE 4.** Comparison of trigger factors frequencies among migraine patients without aura and with aura

| Triggering factors   | Headache type                  | p value |
|----------------------|--------------------------------|---------|
|                      | Migraine without aura (%)     | Migraine with aura (%) |
| Fatty foods          | 10(5.9)                        | 4(4.0)  | 0.501  |
| Chocolate            | 9(5.3)                         | 3(3.0)  | 0.0.33 |
| Cheese               | 31(18)                         | 12(12.0)| 0.0.17 |
| Nuts                 | 6(3.5)                         | 4(4.0)  | 0.84   |
| Watermelon           | 4(2.4)                         | 5(5.0)  | 0.24   |
| Citrus fruits        | 8(4.7)                         | 2(2.0)  | 0.38   |
| Tomato               | 5(2.9)                         | 3(3.0)  | 0.97   |
| Fish                 | 14(8.2)                        | 8(8.0)  | 0.94   |
| Sausage              | 22(12.9)                       | 10(10.0)| 0.43   |
| Pepper               | 60(35.3)                       | 26(26.0)| 0.11   |
| Onion                | 78(45.9)                       | 37(37.0)| 0.15   |
| Tea/Coffee           | 10(5.9)                        | 3(3.0)  | 0.28   |
| Sour food            | 75(44.0)                       | 31(31.0)| 0.33   |
| Smoking              | 94(55.0)                       | 52(52.0)| 0.60   |
| Stress/anxiety       | 130(76.0)                      | 83(83.3)| 0.20   |
| Excitement           | 108(63.5)                      | 67(67.0)| 0.56   |
| Sleep changes        | 114(67.1)                      | 74(74.0)| 0.23   |
| Light                | 103(60.6)                      | 53(53.0)| 0.22   |
| Physical factors     | 102(60.0)                      | 73(73.0)| 0.03   |
| Weather changes      | 98(42.4)                       | 33(33.0)| 0.000  |
| Tiredness            | 108(63.5)                      | 41(41.0)| 0.000  |
| Humidity             | 19(11.2)                       | 10(10.0)| 0.72   |
| Irregular meals      | 46(27.1)                       | 18(18/0)| 0.09   |
| Strong smells        | 72(42.4)                       | 35(35.0)| 0.47   |
| Sexual activity      | 46(27.1)                       | 17(17.0)| 0.05   |
| Menstruation         | 88(55.7)                       | 55(60.4)| 0.05   |

P < 0.05 was considered significant.
DISCUSSION

Migraine, as the most common brain disorder, appears to be associated with neurological and systemic symptoms. Despite a lack of knowledge on migraine’s pathogenesis, a great deal of evidence suggests both genetic and environmental factors, such as hormonal fluctuations, fatigue, stress, meteorological changes, and diet, modulating the threshold of migraine that precedes and evokes a migraine attack (23,24).

**TABLE 5. Logistic regression analyses of trigger factors of migraine patients**

| Triggering factors | migraine | p value |
|--------------------|----------|---------|
|                    | Non-adjusted OR (95%CI) | Adjusted OR (95%CI) |     |
| Fatty foods        | 0.667(0.203-2.184) | 0.924(0.228-3.751) | 0.912 |
| Chocolate          | 0.553(0.146-2.093) | 0.554(0.128-2.393) | 0.554 |
| Cheese             | 0.611(0.298-1.253) | 0.785(0.347-1.773) | 0.560 |
| Nuts               | 1.139(0.314-4.137) | 1.485(0.338-6.514) | 0.600 |
| Watermelon         | 2.184(0.573-8.331) | 2.352(0.563-9.827) | 0.241 |
| Citrus fruits      | 0.365(0.077-1.725) | 0.499(0.088-2.832) | 0.432 |
| Tomato             | 1.021(0.239-4.365) | 1.494(0.266-8.401) | 0.649 |
| Fish               | 0.969(0.392-2.398) | 1.161(0.402-3.353) | 0.783 |
| Sausage            | 0.665(0.294-1.508) | 0.883(0.332-2.350) | 0.803 |
| Pepper             | 0.644(0.373-1.113) | 0.818(0.411-1.629) | 0.568 |
| Onion              | 0.693(0.418-1.149) | 0.881(0.461-1.684) | 0.702 |
| Tea/Coffee         | 0.495(0.133-1.842) | 0.605(0.135-2.701) | 0.510 |
| Sour food          | 0.569(0.338-0.958) | 0.681(0.363-1.278) | 0.231 |
| Smoking            | 0.876(0.534-1.437) | 1.009(0.566-1.800) | 0.975 |
| Stress/anxiety     | 1.502(0.799-2.823) | 1.723(0.849-3.497) | 0.132 |
| Excitement         | 1.166(0.692-1.962) | 1.269(0.690-2.332) | 0.444 |
| Sleep changes      | 1.398(0.807-2.422) | 1.598(0.873-2.923) | 0.129 |
| Light              | 0.734(0.445-1.208) | 0.692(0.387-1.236) | 0.213 |
| Physical factors   | 0.744(0.440-1.258) | 0.850(0.455-1.588) | 0.611 |
| Weather changes    | 0.852(0.519-1.397) | 0.837(0.484-1.448) | 0.525 |
| Tiredness          | 1.098(0.667-1.806) | 1.099(0.626-1.928) | 0.743 |
| Humidity           | 1.438(0.661-3.130) | 1.797(0.746-4.329) | 0.192 |
| Irregular meals    | 0.785(0.434-1.420) | 0.830(0.425-1.618) | 0.584 |
| Strong smells      | 0.766(0.460-1.274) | 0.754(0.424-1.339) | 0.335 |
| Sexual activity    | 0.598(0.242-1.478) | 0.660(0.243-1.794) | 0.415 |

**FIGURE 2. A comparison of common triggers factors of headache with gender (in %).**

*Logistic regression model, non-adjusted.
Logistic regression model, adjusted.
P < 0.05 was considered significant.*
Following an investigation of the related factors of migraine headaches in an Iranian population, it was shown that the incidence of migraine was higher in women than in men, suggesting that migraines in women may be related to the role of female hormones in the development of migraine headaches. It has been reported that with decreasing estrogen levels migraine attacks increase and vice versa (25). Menstruation is the most critical risk factor for the occurrence and persistence of headaches and migraines, with nearly 50% of women reporting menstruation as a potential migraine trigger. In the present study, hormonal triggers such as menstruation affected individuals differently in patients with aura and without aura, indicating that that migraine headache is a fairly common physical condition in Iran, negatively affecting life quality. Among migraine patients, 37% were diagnosed as having migraine with aura, with approximately 50% of cases showing a close relatedness of family history. Reports on migraine related to family history include those in Africa with 14.6%, and among dental students in India with a reported rate of 20.1% (26,27).

Significant association in terms of gender and the presence of a family history of headache and the incidence of migraine in female and male students has also been reported by others (26). The increase in migraine prevalence could be due to stress and unhealthy lifestyle habits such as low physical activity (28). Various studies have shown that the risk of developing headaches is greater in individuals with a family history, smoking habit, sleeping problems, premenstrual period, stressful lifestyle, weather changes, menstruation, hunger etc. (16,29).

In the present research, the most common and most important cause of headache included stress, sleep changes and excitement followed by bright lights, tiredness, and weather change disturbance. Most reports have demonstrated stressful situations as the main trigger of a headache attack (16,30). The present study also shows stressful life events as the commonest trigger in both genders and among migraine subtypes. Considering the relation between sleep changes and headache, Kelman group showed that a relatively shorter sleep duration would lead to increased severity and frequency of migraine attacks (31). At the same time, our results demonstrated an association between physical activity and fatigue in recurrent headaches without aura and aura. Conservative management of both factors may be more effective in preventing headaches (32).

Nearly one-half of the subjects with migraine (44.65%) in the present study believed that spending a long time outside under the sunlight and being exposed to extreme heat made their headaches worse, indicating the role of weather changes as trigger factors, as demonstrated by others (16,33). Although the effect of food factors intensifying migraine headache has been reported by some authors (34), no such relation was detected in the present study (16). Such discrepancies may be explained by existing differences between population’s social habits and culture. Investigating the role of diet factors in migraines can be complicated, as various triggers and variables may alter the pain threshold in the individual (17). Dietary intervention in disease contains many components that may interact with various genes in several different ways. If any, interventions to modify targeted diet may lead to the replacement of a proper diet with medication (23). Avoiding foods that are effective in the intensification of migraine headaches could help relieve pain, headache duration, and the number of migraine attacks (22). Considering that by reducing stress in the workplace and stressful life events, one can relieve migraine headaches to some extent, it is thus possible to reduce the frequency and severity of migraine attacks by informing migraine patients of the factors affecting their occurrence migraine attacks.

**CONCLUSIONS**

In conclusion, the epidemiological data on the type and quality of headache, family history, related factors and intensifying factors of migraine headaches were obtained. Socio-economic impacts of migraine on the quality of life of patients and society are significant. To improve the health of the community, strategies for managing these factors should be effectively implemented. Also, studies on the prevention, reduction, and control of factors affecting migraine headaches with regard to reinforcing factors can help improve the health of the communi-
ty. Considering the contradictory reports on the involvement of different migraine trigger factors, it is necessary to conduct studies aimed at defining the effect of trigger factors on the process of migraines in a more accurate manner.

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