# Evaluation of Patent Foramen Ovale in Migraine with aura Headaches

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### **Abstract**

**Introduction**:Migraine is one of the most common headaches, and is one of the most common disabling medical conditions worldwide. Migraine has two types, with aura and without aura. Nearly 15% of migraines are with aura, and the most common aura is visual. Patent Foramen Ovale (PFO) is the most common defect in atrial septum of heart, and prevalence rate population is 25%. Lately PFO has been recognized to be a risk factor for migraine with aura.

**Methods and mareials**: This study is cross sectional and has been done on 50 patients of migraine with aura, and 25 normal people (without migraine) as control group. We used saline agitated test by Doppler sonograph for recognizing PFO.

**Results**: The rate of PFO in patients' group was 44% and in control group was 20%. The rate of PFO is more in patients with rate of headaches more than 3 times in a month.

**Conclusion**: migraine with aura has relation with PFO, especially for patients with frequent attacks more than 3 times per months. The closure of PFO in these patients need to be studied more, especially that there are feasible procedures available now.

Keywords: migraine, migraine with aura, PFO, patent foramen oval, saline agitated test.

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#### Introduction

Of all the painful states that afflict humans, headache is undoubtedly the most frequent and rivals backache as the most common reason for seeking medical help.<sup>(1)</sup>

The lifetime prevalence for any type of headache as estimated from populationbased studies is more than 90% for male and 95% for female. Migraine is one of the most common headaches. Prevalence rate for migraine in USA is 6.5% for male and 18.2% for female, and can begin at any age, but most common in adolescence. Migraine is one of the most common disabling medical conditions worldwide. Migraine is classified into two general groups (by International Headache Society) with aura and without aura, and other types or subtypes that are not frequent. Nearly 15% of migraines are with aura, and the most common aura is visual. (2)

In a study done in Iran, the prevalence rate is 7% for male, and 14.2% for female. (3)

Patent foramen ovale (PFO) is the most common defect in atrial septum of heart, and prevalence rate population is 25%. (2) This defect often has no clinical signs or symptoms, and remains undiagnosed. But it is recognized as a factor of right to left shunt in heart. (4) Lately PFO has been shown to be a factor for producing migraine with aura and stroke in voung adults. it's considered. probably with nonischemic mechanisms of migraine. (5) However, there is currently insufficient evidence to support a causal link

between PFO and migraine. (6) PFO can be diagnosed by transesophageal echocardiogram which is an invasive procedure. Another method that can be use for PFO diagnosis is agitated saline test which is an easy and non invasive procedure. This method is easily tolerated by patients and needs Doppler ultrasonography to detect echogenous bubble of air. This test has 94% sensitivity and 97% specificity. (7) In an evaluation done, the rate of PFO in migraine without aura is the same as normal population, but in migraine with aura is 62%. (8) We didn't find any study about frequency of headaches and duration of disease. In this study we evaluated frequency of PFO in migraine with aura, control group (without migraine) and additionally, relation of frequency of PFO to frequency of attack of headaches per month and duration of disease.

#### Methods and materials

This study was cross sectional and has been done on 50 patients of migraine with aura, and 25 normal people (without migraine) as control group. Patients were selected from admitted migraine patients to Shafa medical center. The goal and method of exam was described for the patients. For every patient a form including sex, age, age of onset of headaches, type of headache, unilateral, bilateral, duration of attacks, familial history of migraine, precipitating factors like specific foods, stress, menstrual periods, frequency of attacks in a months, and signs or symptoms associated with headaches,

nausea, vomiting, photophobia, phonophobia was filled. The control group consisted of normal population without history of migraine headache and cardiovascular disorders.

Inclusion criteria are:

- 1- migraine with aura (IHS criteria) diagnosed by neurologist. (9)
- 2- Normal physical examination, specially cardiopulmonary diseases has been ruled out.
- 3- Signed form of information by the patient.

The test has been done by a Doppler sonograph Pulse Wave (PW) instrument made in Germany and a software box (DWL2.5X2) to detect blood flow with 2 mega hertz's probe with gain 50, deep 450-55 mm fixed on right Middle Cerebral Artery (MCA) from temporal window.

Saline agitated test: The patient lay on the bed, and an angio-catheter No 20 use to put in brachial vein and then joint with special connector.

9 ml of saline normal solution, 0.5 ml air and 0.5 ml blood of patient shake well to produce a homogenize solution. This solution inject with 1 ml/second speed in every phase, but may repeat to three phases. In first phase without Valsalva maneuver, if bubble of micro emboli was detected in MCA by PW when homogenize solution is injected, test is complete. If bubble of micro emboli has not been detected, second phase should be done. This phase is the same as first phase but the patients has deep inspiration and after that sever expiration (equivalent of Valsalva maneuver), this maneuver increase right to left shunt of heart, and as a result the micro emboli of air passes from shunt and enter to MCA, and then can be detected by PW instrument. (10) If micro emboli was not detected, the third phase will be done. Third phase is the same as other phases, but 5 minutes after expiration ending has been done. On monitor of PW instrument the waves were evaluated and findings were detected on form.

#### Results

50 patients with migraine with aura were evaluated (78% female and 22% male) with mean age of 37.54 years. 25 normal people were studied.

Table 1 shows the rate of frequency of PFO in patients and control groups (male and female). Mean age of control group was 35.77 years old (without significant difference with patients' group).

Table 2 shows the frequency of headaches per month in patients and the rate of PFO.

The average duration of disease is 8.43 years without significant sex difference and also between PFO positive and PFO negative. Headache frequency is significantly different groups. Other variables, like foods, age, onset-age of headaches. type headache, unilateral, bilateral, duration of attacks, familial history of migraine, precipitating factors such as special foods, stress, menstrual periods, and signs or symptoms associated with headaches. nausea, vomiting, photophobia, phonophobia were not statistically different.

#### **Discussion**

Studies linking migraine with aura and a hole in one's heart have been big news in recent years. The hole is present in fetuses to allow blood to pass from one side of heart to another. In most people, the gap will be closed at or after birth. For an estimated 25% of the US population, it doesn't close completely and PFO forms. But in other studies the rate of PFO different, A 10% prevalence rate in ultrasound evaluation, and 27% autopsy was reported, but this rate is equal in men and women. (11) In a study PFO was found in 9.2% of all patients and also seen in all age groups. Incidence in patients with ages of 40-49 years was greater than aged 70-79 vears (12.86% vs 6.15%). (12) In a study, the rate of PFO in migraine with aura was reported to be 41%. (13) In another study, this rate was 48% in migraine with aura and 23% in migraine without aura<sup>(8)</sup> In a study which has been done in Iran, this rate for migraine with aura was 38.5% and for migraine without aura was 26.7%. (14) In a study which has been done by transesophageal echocardiography, the rate of PFO was 47%, (15) and another study the rate of PFO is twice in migraine with aura. (16) About relation of migraine with aura PFO, there not and is definite mechanism, genetically be may association between migraine with aura and PFO.<sup>(17)</sup> patients In with paradoxical cerebral embolism. migraine headaches are more frequent than in the general population, and transcatheter closure of PFO, results in

complete resolution marked or reduction in frequency of migraine headaches. (18-19) In a study, rate of PFO in migraine with aura is  $62\%^{(20)}$  In our study the rate of PFO in migraine with aura is 44%, and 20% in control group (P=0.005), but without significant sex difference, for the patients that have transesophageal the PFO. echocardiogram has been done, and results for all patients was positive. In our study those patients who had more frequent headaches, the **PFO** frequencies is also more.

That's important, because PFO may be one of the causes of strokes in young. The frequency of PFO in stroke patients is higher than among the general population, particularly individuals with stroke of unknown etiology (cryptogenic stroke) and in younger patients. (21) Wildly speculated that people have PFO, die earlier than those who don't have. (22) Accordingly individuals with a history of migraine with aura should be screened for the presence of a large shunt before they dive or exposed to subatmospheric decompression. (23) Migraine may be associated with other disorders, example depression. (24) Our study proved other studies about association of migraine with aura and PFO. The PFO has association with stroke, thus we must consider these patients. because several secondary stroke have prevention strategies been developed: antiplatelet agents, oral anticoagulants, percutaneous closure, surgical repair, (6-25) and thermal PFO closure is feasible, safe, effective and

may be possible in future. (26)

Migraine with aura has relation with PFO, especially in patients with frequent attacks more than 4 times per months. The closure of PFO in these patients need to be studied more, especially that, there are feasible procedures available now.



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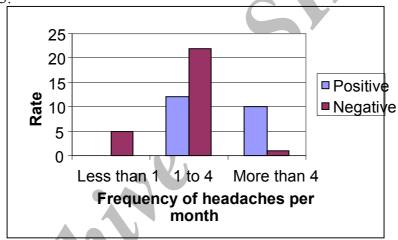
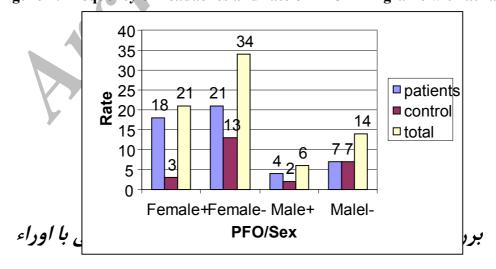


Figure 2: Frequency of headaches and rate of PFO in migraine with aura



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## چکیده

سابقه وهدف: سر درد میگرنی یکی از شایع ترین سر دردهاست، و یکی از شایع ترین عوامل ناتوان کننده پزشکی در دنیاست. سر درد میگرنی بر دو نوع است: با اوراء و بدون اوراء، تقریبا ۱۵٪ سر دردهای میگرنی با اوراء هستند و شایع ترین اوراء بینائی است.

باز بودن سوراخ بیضی شایع ترین نقص بین دیواره ای دهلیزی قلب است و ۲۵٪ افراد جامعه سوراخ بیضی باز دارند. سوراخ بیضی باز اخیرا بعنوان یکی از عوامل ایجاد کننده سر درد میگرن با اوراء شناخته شده است. روش بررسی: این مطالعه مقطعی روی ۵۰ بیار با سر درد میگرن با اوراء و ۲۰ فرد سالم بدون سر درد میگرن انجام شد. با استفاده از تست سالین آژیته و با کمک از دستگاه سونوگرافی داپلر تشخیص انجام شد. یافته ها : میزان فراوانی سوراخ بیضی باز در گروه بیماران سردرد میگرنی با اوراء ۴۴٪ و در گروه شاهد ۲۲٪ بود. بیمارانی که بیش از سه بار سردرد میگرنی در یک ماه داشتند میزان شیوع سوراخ بیضی باز بیشتری داشتند

**نتیجه گیری:** سردرد میگرنی با اوراء با سوراخ بیضی باز ارتباط دارد، بخصوص در بیمارانی که بیش از سه بار در یک ماه سردرد دارند. بستن سوراخ بیضی باز با به در دسترس بودن امکانات باید مورد توجه بیشتر قرار گیرد.

واژگان کلیدی: سر درد میگرنی، سر درد میگرنی با اوراء، سوراخ بیضی باز و تست سالین آژیته.