

Figure 1. A: cumulative survival. B: function of cumulative incidence of recurrence.

the need to perform additional investigations at the time of CM diagnosis to look for other concurrent cancers and open avenues of clinical research into the etiology and natural history of this type of tumor.

In conclusion, the most precise data on CM incidence show an increase in the diagnosis of these tumors in recent years and demonstrate the essential role of echocardiography.

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## Psychiatric Symptoms and Sex-related Differences in Patients With Myocardial Infarction With Nonobstructive Coronary Arteries



### Síntomas siquiátricos y diferencias relacionadas con el sexo en pacientes con infarto de miocardio con arterias coronarias no obstructivas

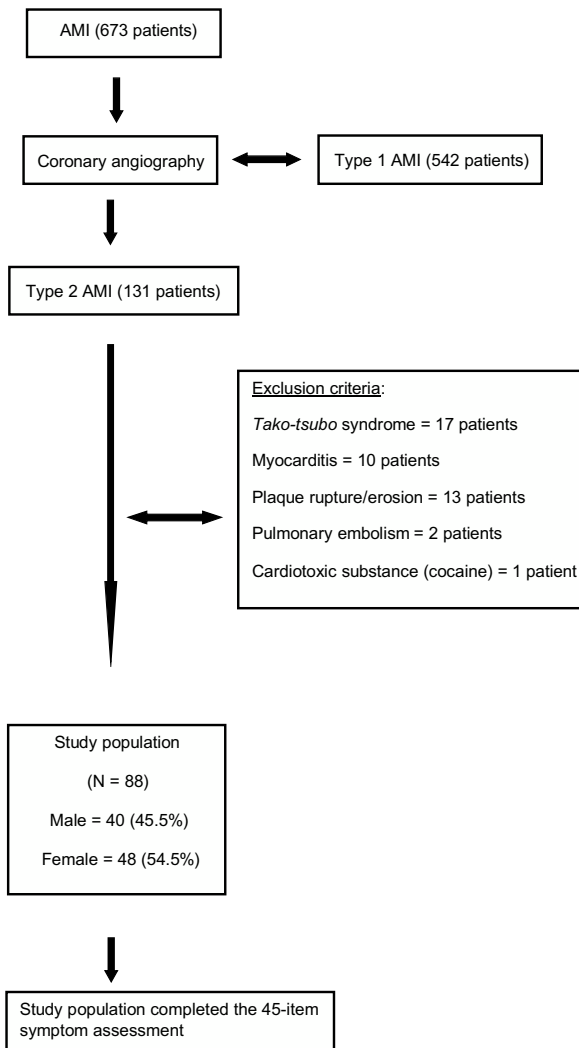
#### To the Editor,

In a recent Position Paper of the European Society of Cardiology,<sup>1</sup> the following criteria for diagnosis of myocardial infarction with nonobstructive coronary arteries (MINOCA) were proposed: a) acute myocardial infarction according to the criteria defined by the III universal definition; b) absence of  $\geq 50\%$  stenosis on angiography; c) exclusion of other clinically overt specific etiologies. Anxiety and mood disorders seem to be more common in women than in men, and there is emerging evidence linking anxiety to coronary artery disease (CAD) development, particularly among women.<sup>2</sup> A previous study demonstrated sex differences in the prevalence of psychiatric disorders in patients with unstable angina.<sup>3</sup> Little is known about sex differences in the prevalence of

psychiatric symptoms in patients with MINOCA. The aim of this study was to examine the relationship between sex and psychiatric symptoms in patients with MINOCA.

We prospectively evaluated 131 patients with a final etiologic diagnosis of MINOCA who underwent coronary angiography at the Cardiology Department of a University Hospital from October 1, 2011 to December 31, 2017. Nonobstructive CAD was defined as the presence of coronary stenosis  $> 0\%$  but  $< 50\%$  of lumen diameter in at least 1 major epicardial coronary artery.<sup>1</sup> We excluded 17 patients with a diagnosis of *tako-tsubo* syndrome confirmed by echocardiography or cardiac magnetic resonance, 10 patients with a suspected diagnosis of myocarditis confirmed by cardiac magnetic resonance, 13 patients without obstructive CAD but with evidence of coronary thrombosis on an unstable plaque confirmed by intravascular ultrasound, 2 patients with coronary embolism, and 1 patient who underwent cardiotoxic substance administration (cocaine). Hence, 88 patients were included in the study (Figure 1). The study was approved by the local ethics committee and all patients gave written informed consent before angiography.

Once their clinical condition had stabilized, all patients completed the 45-item Symptom Assessment (SA-45).<sup>4</sup> The SA-



**Figure 1.** Study flowchart. The study cohort. AMI, acute myocardial infarction.

45 assesses 9 psychopathological domains: hostility, somatization, depression, anxiety, interpersonal sensitivity, obsession-compulsion, phobic anxiety, paranoid ideation, and psychoticism. Each domain contains 5 items that are scored on a 5-point scale of distress (0, none; 1, a little; 2, moderate; 3, quite a bit; and 4, extreme); the total score reflects the severity of the corresponding psychiatric domain<sup>4</sup> (Table 1). In addition, we collected the following variables for the study: age, sex, body mass index, cardiovascular risk factors, left ventricular ejection fraction, psychiatric history in first-degree family members, education level, marital status, and treatment at discharge. The relationship between sex and psychiatric symptoms was analyzed by means of logistic regression. The SPSS v20 was used for all calculations.

The mean patient age was 53 years, and 45.5% were men. There were no significant differences in cardiovascular risk factors, previous psychiatric history in first-degree family members, education level, or marital status between men and women. Analysis of the psychiatric symptoms showed that women had higher scores than men in somatization and phobic anxiety (Table 1). After adjustment by baseline characteristics, a multivariable logistic regression model showed that the sex difference was statistically significant with a female-male odds ratio for phobic anxiety of 1.2 (95%CI, 1.04-1.40; *P* = .01).

The main and original finding in our study assessing sex differences in psychiatric symptoms among MINOCA patients was

**Table 1**

SA-45 Questionnaire Results of the Study Population and Baseline Characteristics Between Male and Female Patients With MINOCA

SA-45 Questionnaire results in 88 consecutive patients with MINOCA			
Anxiety			7.9 ± 3.8
Depression			5.6 ± 4.4
Interpersonal sensitivity			5.2 ± 3.8
Hostility			3.9 ± 3.8
Obsessive-Compulsive			6.1 ± 3.9
Psychoticism			2.8 ± 2.7
Paranoid ideation			8 ± 4.3
Somatization			4.4 ± 3.3
Phobic anxiety			3.5 ± 3.5
Baseline characteristics of 88 consecutive patients with MINOCA: comparison between male and female patients			
Variables	Men (n = 40)	Women (n = 48)	<i>P</i>
Age, y	53.8 ± 8.5	52.9 ± 8.4	.6
Body mass index, kg/m <sup>2</sup>	27.8 ± 3.3	27.6 ± 3.7	.8
Hypertension	22 (55)	27 (56.2)	.9
Smoker	20 (50)	23 (47.9)	.8
Dyslipidemia	26 (65)	25 (52.1)	.22
Diabetes mellitus	12 (30)	14 (29.2)	.93
Left ventricular ejection fraction, %	55.8 ± 9.4	58.1 ± 9.5	.25
Higher education	9 (22.5)	12 (25)	.8
Previous psychiatric disease	3 (7.5)	5 (10.4)	.7
Civil status, married	26 (65)	36 (75)	.4
Time to perform the CC, h	30.3 ± 11.4	32.7 ± 11.1	.31
SA-45 Questionnaire			
Depression	5.2 ± 3.9	5.9 ± 4.8	.5
Anxiety	6.8 ± 3.4	8.8 ± 3.9	.15
Interpersonal sensitivity	5.3 ± 3.3	5.2 ± 4.2	.9
Hostility	3.5 ± 3.4	4.3 ± 4	.3
Obsessive-compulsive	6.4 ± 3.4	5.8 ± 4.3	.5
Psychoticism	2.6 ± 2.3	3 ± 3	.6
Paranoid ideation	7.7 ± 3.5	8.2 ± 4.9	.6
Somatization	3.4 ± 2.9	5.2 ± 3.4	.009
Phobic anxiety	2.4 ± 3.5	4.5 ± 3.3	.004
In-hospital medication			
Antiplatelets	38 (95)	48 (100)	.11
Statins	40 (100)	48 (100)	*
Beta blockers	17 (42.5)	20 (41.7)	.9
Calcium channel antagonists	5 (12.5)	9 (18.8)	.6
ACEI/angiotensin II antagonist	11 (27.5)	21 (43.8)	.13
LMWH	40 (100)	48 (100)	*
Vasodilators	36 (90)	45 (93.8)	.51
Treatment at discharge			
Aspirin	31 (77.5)	45 (93.8)	.03
Beta-blockers	17 (42.5)	20 (41.7)	.9
Calcium channel antagonists	5 (12.5)	9 (18.8)	.6
ACEI/angiotensin II antagonist	11 (27.5)	21 (43.8)	.13
Statins	18 (45)	30 (62.5)	.22

ACEI, angiotensin-converting enzyme inhibitor; CC, cardiac catheterization; LMWH, low molecular weight heparin; SA-45, Symptom Assessment-45. Data are presented as No. (%) or mean ± standard deviation.

\* Statistic is not calculated for this variable because it is a constant.

that female patients had higher phobic anxiety than male patients. Anxiety disorders are the most prevalent mental disorders and are associated with immense health care costs and a high disease burden. According to large population-based surveys, up to 33.7%

of the population is affected by an anxiety disorder during their lifetime.<sup>2</sup> Phobic anxiety, characterized by an unreasonable fear when exposed to specific situations such as enclosed spaces, heights, or crowds, is the predominant complaint in approximately half of these individuals.<sup>2</sup>

In our study, female MINOCA patients showed significantly higher scores for phobic anxiety than male MINOCA patients. A similar sex distribution was seen in a prior study, in relation to elevated phobic anxiety and female CAD patients.<sup>5</sup> Watkins et al.<sup>5</sup> reported that phobic anxiety levels were high in women with CAD and may be a risk factor for cardiac-related mortality in women diagnosed with CAD. In that prospective cohort study, 947 CAD patients were included, and participants completed the phobic anxiety subscale of the Middlesex Hospital Questionnaire. CAD was defined as  $\geq 75\%$  occlusion of 1 coronary artery. Female CAD patients reported significantly elevated phobic anxiety levels compared with male patients ( $P < .001$ ). In women, phobic anxiety was associated with a 1.6-fold increased risk of cardiac mortality and a 2.0-fold increased risk of sudden cardiac death but was not associated with increased mortality risk in men. As in our study, Watkins et al.<sup>5</sup> found an association between sex and phobic anxiety in patients with CAD. Their population was comparable to ours except that the patients at the time of enrollment had unstable or stable angina and the presence of  $\geq 75\%$  stenosis on angiography. In our study, we enrolled only MINOCA patients. Our study has some limitations. First, this is a single-center study. Second, there was no control group. Third, the study population was not large.

Phobic anxiety should be tested in female MINOCA patients, as they reveal novel targets for the development of novel pharmacotherapies that could be specifically tailored to the physiology of women.<sup>6</sup>

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## Atrial Septostomy in Children With Pulmonary Hypertension



### Septostomía auricular en niños con hipertensión pulmonar

#### To the Editor,

Pulmonary hypertension (PH) is an irreversible progressive disease that, unless diagnosed early, can lead to right ventricular failure and end-stage heart failure (HF). Significant advances have been made in pharmacological treatment,<sup>1</sup> but life expectancy remains short—57% at 5 years—and the disease is more aggressive in children than in adults.<sup>2</sup>

Atrial septostomy (AS) is indicated in children with PH in functional class III or IV of the Ross classification who experience recurrent syncope and/or right HF refractory to medical treatment, as a bridge to lung transplant.<sup>3,4</sup> The main contraindications

are a mean atrial pressure  $> 20$  mmHg and oxygen saturation  $< 90\%$ .<sup>5</sup>

AS can be performed using a number of techniques; one of the more commonly used, and which is the practice in our hospital, involves placing a stent in the interatrial septum. This technique is performed with a 7-Fr sheath and a pediatric Brockenbrough needle, which passes first through the inferior vena cava in a 15-mm snare that is positioned from the contralateral vein. Once the right atrial pressure has been checked to ensure it does not contraindicate the procedure, the septum is punctured, the sheath is positioned in the left atrium, and a coronary guidewire is advanced to the pulmonary vein. A 10 × 19 mm Palmaz Genesis large stent is positioned in the center of the septum. The snare is then positioned halfway along the stent, and the stent is dilated, restricted in the middle by the snare so that it takes on an hourglass shape and becomes highly stable. Balloon dilatation is then performed with coronary or peripheral balloons, based on