

## Article

# Association of occupational distress and low sleep quality with syncope, presyncope and falls in workers

Nicola Magnavita <sup>1,2,3,\*</sup>, Reparata Rosa Di Prinzio <sup>1</sup>, Gabriele Arnesano <sup>1</sup>, Anna Cerrina <sup>3</sup>, Maddalena Gabriele <sup>3</sup>, Sergio Garbarino <sup>1,4</sup>, Martina Gasbarri <sup>3</sup>, Angela Iuliano <sup>1</sup>, Marcella Labella <sup>3</sup>, Carmela Matera<sup>3</sup>, Igor Mauro <sup>1</sup>, Franca Barbic <sup>1,5,6</sup>

<sup>1</sup> Postgraduate School of Occupational Health, Università Cattolica del Sacro Cuore, Rome, Italy; [repdip@gmail.com](mailto:repdip@gmail.com) (R.R.D.P.), [gabrielearnesano93@gmail.com](mailto:gabrielearnesano93@gmail.com) (G.A.), [iuliano\\_angela@yahoo.it](mailto:iuliano_angela@yahoo.it) (A.I.), [mau.igor91@yahoo.it](mailto:mau.igor91@yahoo.it) (I.M.)

<sup>2</sup> Department of Woman, Child & Public Health, Fondazione Policlinico Universitario A. Gemelli IRCCS, Rome, Italy.

<sup>3</sup> Local Sanitary Unit Roma4, Civitavecchia, Italy; [anna.cerrina@aslroma4.it](mailto:anna.cerrina@aslroma4.it) (A.C.), [maddalena.gabriele@aslroma4.it](mailto:maddalena.gabriele@aslroma4.it) (M.Gab.), [martina.gasbarri@aslroma4.it](mailto:martina.gasbarri@aslroma4.it) (M.Gas.), [marcella.labella@aslroma4.it](mailto:marcella.labella@aslroma4.it) (M.L.), [carmela.matera@aslroma4.it](mailto:carmela.matera@aslroma4.it) (C.M.),

<sup>4</sup> Department of Neuroscience, Rehabilitation, Ophthalmology, Genetics and Maternal-Infantile Sciences (DINOEMI), Genoa, Italy [sgarbarino.neuro@gmail.com](mailto:sgarbarino.neuro@gmail.com) (S.G.),

<sup>5</sup> Department of Biomedical Sciences, Humanitas University, Milan, Italy; [franca.barbic@hunimed.eu](mailto:franca.barbic@hunimed.eu) (F.B.)

<sup>6</sup> Internal Medicine, IRCCS Humanitas Research Hospital, Rozzano-Milan, Italy.

\* Correspondence: Nicola Magnavita, [nicolamagnavita@gmail.com](mailto:nicolamagnavita@gmail.com) tel.: +39 3473300367

**Abstract:** Syncope and presyncope occurring during work can affect safety and impair occupational performance. Few data are available regarding the prevalence of these events among workers. The possible role of sleep quality, mental stress and metabolic disorders in promoting syncope, presyncope and falls in workers is unknown. In the present study, 741 workers (male 35.4%; mean age 47±11 years), employed in different companies, underwent clinical evaluation and blood tests and completed questionnaires to assess sleep quality, occupational distress and mental disorders. The occurrence of syncope, presyncope and unexplained falls during their working life was assessed by an ad hoc interview. The prevalence of syncope, presyncope and falls of unknown origin was 13.9%, 27.0%, and 10.3%, respectively. The occurrence of syncope was associated with an increased risk of occupational distress (adjusted Odds Ratio aOR: 1.62, Confidence Intervals at 95%: 1.05-2.52), low sleep quality (aOR: 1.79 CI 95%: 1.16-2.77) and poor mental health (aOR: 2.43 CI 95%: 1.52-3.87). Presyncope was strongly associated with occupational distress (aOR: 1.77 CI 95%: 1.25-2.49), low sleep quality (aOR: 2.95 CI 95%: 2.08-4.18) and poor mental health (aOR: 2.61 CI 95%: 1.78-3.84), while no significant relationship was found between syncope or presyncope and metabolic syndrome. These results suggest that occupational health promotion interventions aimed at improving sleep quality, reducing stressors and increasing worker resilience might reduce syncope and presyncope events in the working population.

**Keywords:** Loss of consciousness; Mental health; Working life; Effort Reward Imbalance; Sleep Disorders; Health promotion; Workplace

## 1. Introduction

Syncope, which is defined as a transient loss of consciousness due to cerebral hypoperfusion, is characterized by a rapid onset, short duration and complete, spontaneous recovery [1, 2]. It can be the result of a reduction in cardiac output caused by serious cardiovascular disorders (5-10%) such as arrhythmias or structural heart disease or is more commonly due to neuro-mediated mechanisms such as vasovagal syncope or syncope associated with orthostatic hypotension [1-3]. Interestingly, prolonged bed rest in various

clinical conditions (e.g., after surgery or trauma) may increase the risk of syncope [4]. Syncope is also quite frequently associated with constitutional hypotension and often underestimated in young women [5].

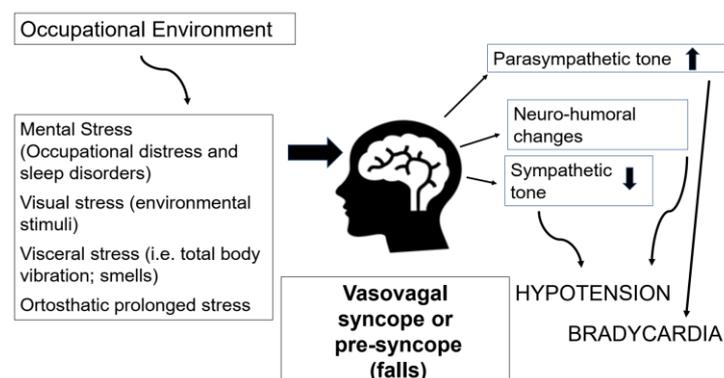
A transient loss of consciousness (T-LOC) constitutes a potential hazard if it occurs in the workplace. More than 90% of T-LOCs are due to syncope, epileptic seizures or psychogenic non-epileptic seizures [2, 6, 7]. A prodromal period characterized by symptoms such as light-headedness, nausea, sweating, weakness, and visual disturbances may indicate that syncope is imminent and is defined as “presyncope” [8]. Lastly, unexplained falls may hide episodes of syncope or presyncope [9].

Recently, the management of workers suffering from syncope has aroused increasing interest [2, 10-12] due to the potentially fatal outcome for the individual or third parties when an episode occurs in a worker undertaking hazardous tasks. In a Danish nationwide cohort, syncope was associated with a 1.4-fold greater risk of occupational accidents and a 2-fold higher risk of termination of employment compared with the general population workforce [13]. The incidence of syncope in the military has been estimated at 7.2 cases per 1,000 person-years [14]. A Polish study found that 4.7% and 14.8% of operating room staff had experienced at least one episode of syncope and presyncope, respectively [15].

The occurrence of these problems is rarely reported to health services. The Framingham offspring study revealed that 44% of the participants with an episode of loss of consciousness failed to undergo medical evaluation [16], and a much higher percentage was observed in the younger population. In the Netherlands, admission to the Emergency Department (ED) accounts for only about 1% of expected syncope episodes in the general population, thus indicating that syncope and presyncope are underestimated [17].

In EDs, the management of patients suffering from T-LOC (syncope included) presents a challenge [1, 2, 10], since the emergency physician must first of all exclude the most serious causes of syncope before discharging a patient. However, patients diagnosed with benign vasovagal syncope (the most common type of syncope) often receive no explanation of the event, leaving them with a consequent fear of syncope recurrence and a possible reduction in the quality of life.

Syncope and presyncope are common during working life [12, 18-20] and may be triggered by occupational tasks requiring prolonged standing, exposure to a hot climate or frequent changes of posture [11]. Furthermore, both central and peripheral initiating factors such as mental, visceral, visual, and orthostatic distress, potentially present in work environments or during specific occupational tasks, may facilitate vasovagal syncope [21], particularly in hypersusceptible individuals. Environmental stressors, including psychosocial stress, may also play a role in promoting vasovagal syncope, which is the most common cause of syncope, particularly during working life, by facilitating the mechanisms suggested by Mosqueda-Garcia [22]. Figure 1 illustrates the mechanisms activating vasovagal syncope and the possible role of the occupational environment.



**Figure 1.** Potential role of environmental stressors in promoting vasovagal syncope. (from Mosqueda-Garcia [22], modified).

Syncope recurrence is about 22% at 2 years from the syncope index [23]. Interestingly, the highest risk of syncope recurrence in the first 6 months after the syncope index was observed in individuals of working age [12].

The occupational physician, whose aim is to improve workers' health, is also responsible for promptly identifying the symptoms of diseases that may interfere with work safety.

The present study, which was performed during periodic medical examination in the workplace, aimed at quantifying the prevalence of syncope and presyncope occurring during the working life of individuals employed in different companies. We also assessed whether these episodes were associated with low sleep quality, occupational distress and mental health. In addition, an evaluation was made in the same population of a possible association between metabolic syndrome and syncope and presyncope.

## 2. Materials and Methods

### 2.1. Study population

All 754 workers consecutively enrolled for their periodic medical examination from July to December 2020 were asked to fill in an anamnestic questionnaire regarding the occurrence of syncope, presyncope and falls - including those without apparent cause - during their working life. Eight workers refused to participate in the survey, and another 5 provided incomplete answers and were therefore excluded from the study.

All subjects provided informed consent to the processing of personal data for research purposes. The research was conducted in compliance with the principles of the Helsinki Declaration as part of the mandatory medical surveillance of workers in the workplace, approved by the Institutional Review Board and reviewed by the Ethics Committee of the University.

### 2.2. Questionnaire

#### 2.2.1 Prevalence of syncope, presyncope, and falls

To ascertain the prevalence of syncope, presyncope and unexplained falls, the workers were asked to answer the following questions. For syncope: "In your working life have you ever experienced a temporary loss of consciousness, fallen down and then spontaneously recovered consciousness?"; for presyncope: "In your working life have you ever been on the verge of fainting or felt as if you were fainting with cold sweats, intense weakness and the need to sit or lie down?"; for falls of unknown cause: "In your working life have you ever fallen to the ground?" and: "Did the fall occur without a clear accidental reason (tripping, pushing, sliding, etc.)?". The number and date of episodes, the presence of associated trauma, familiarity with similar episodes and the occurrence of episodes during working activity were also assessed. These questions were added to a standard questionnaire used during periodic medical examination.

#### 2.2.2. Occupational stress

Occupational stress was assessed using the Italian version [24, 25] of the Siegrist Effort/ Reward imbalance model [26, 27]. The questionnaire contains 10 questions whose answers are graded according to a 4-point Likert scale. The Effort sub-scale, which determines the psychological effort made to work, consists of 3 questions and the score ranges from 3 to 12. The Reward sub-scale is composed of 7 questions and the score ranges from 7 to 28. The weighted relationship between the two questions, Effort/ Reward imbalance index (ERI) is conventionally considered an expression of distress if higher than one. The reliability of the questionnaire (Cronbach's alpha) in this survey was equal to 0.822 (Effort), and 0.710 (Reward).

#### 2.2.3 Sleep quality

The quality of sleep was assessed using the Italian version [28] of the Pittsburgh questionnaire [29] that consists of 18 questions that form the Pittsburgh Sleep Quality Index (PSQI). An overall score equal to or greater than 5, corresponds to poor sleep quality (bad sleepers). Cronbach's alpha in this study was 0.835.

#### 2.2.4. Mental health

The mental health of workers was assessed using the Italian version [30, 31] of the 12-item General Health Questionnaire (GHQ-12) [32-34], which is a reliable screening instrument for psychological distress and a measure of the common mental health problems/domains of depression, anxiety, somatic symptoms and social withdrawal, rated on a 4-point scale. We used the scoring method (0-0-1-1) with the cut-off level recommended by the authors [35]. A value  $\geq 3$  was classified as Low Mental Health. The reliability of the questionnaire in this study was 0.868.

#### 2.3. Medical examination and blood tests

The occupational physician in charge carried out a clinical evaluation of all the workers participating in the study. In addition, workers' height and weight were recorded to quantify body mass index (BMI) and their waist circumference was measured at the narrowest point between the lower costal (10th rib) and the iliac crest. A sphygmomanometer was used to measure blood pressure while supine and during active standing. Venous fasting blood samples were collected in plain tubes and centrifuged at 3,000 rpm for 10 min at room temperature, and serum samples were frozen at  $-20^{\circ}\text{C}$  until assayed. Fasting blood glucose (FBG) and blood lipid profile (total cholesterol, high-density lipoprotein, triglycerides, and low-density lipoprotein) were determined using an enzymatic assay kit.

##### 2.3.1 Metabolic syndrome prevalence

Components of metabolic syndrome were defined according to the International Diabetes Federation (IDF) [36], the National Cholesterol Education Program Expert Panel on Detection Evaluation and Treatment of High Cholesterol in Adults (NCEP/ATPIII) [37] and the American Association of Clinical Endocrinologists (AACE) [38]. Obesity was defined as BMI  $\geq 25 \text{ kg/m}^2$ , or a waist circumference of  $\geq 94 \text{ cm}$ . for men and  $\geq 80 \text{ cm}$  for women, while hypertriglyceridemia was defined as a serum triglyceride level  $>150 \text{ mg/dL}$  ( $1.7 \text{ mmol/L}$ ). A low level of high-density lipoprotein (HDL) serum cholesterol was defined as a serum HDL-cholesterol  $<40 \text{ mg/dL}$  ( $1.03 \text{ mmol/L}$ ). A systolic blood pressure  $>130 \text{ mmHg}$  and/or a diastolic blood pressure  $>85 \text{ mmHg}$  or drug treatment for hypertension were classified as high blood pressure, while a plasma glucose level  $>100 \text{ mg/dL}$  ( $5.6 \text{ mmol/L}$ ) or the presence of hypoglycaemic drug treatment were classified as high fasting glucose. The presence of three or more abnormalities in the aforementioned components was considered to constitute metabolic syndrome (MetS) [39].

#### 2.4. Statistics

Socio-demographic features were analyzed using frequency or statistical distribution for categorical and continuous variable respectively. In accordance with the literature [40], the age of 55 years was used as a cut-off to divide participants into older and younger workers. The Chi square test was used to compare case distribution by gender and age, while the unpaired Student's t test was used to compare occupational stress, sleep quality, mental health and metabolic syndrome prevalence (treated as continuous variables) in workers with and without syncope, presyncope and unexplained falls.

A logistic regression analysis was performed to ascertain if the occupational variables investigated could predict the occurrence of syncope, presyncope and falls, using socio-demographics as correction factors. The estimated effect was presented in terms of adjusted odds ratio (aOR) and 95% confidence intervals. Statistical analyses were performed using the IBM Statistical Package for Social Sciences, SPSS, version 26.0 statistical software. The significance criterion for a two-tailed P value  $\leq 0.05$  was applied.

### 3. Results

Overall, 741 subjects classified as fit to work (male 262, 35.4%; female 479, 64.6%; mean age  $47 \pm 11$  years) participated in the study, corresponding to 98.3% of the workers selected.

Table 1 shows the prevalence of T-LOC episodes and stress, sleep problems and metabolic syndrome in all workers, according to gender and age. More than 50% of workers reported at least one episode of syncope, presyncope or falls of unknown origin during their period of employment. Syncope and presyncope were more frequent in women than in men. Syncope was reported more frequently in young workers than in those over 55 years of age. Falls were quite frequent, affecting more than one in three workers. Interestingly, unexplained falls resulting from no apparent cause affected 10% of the workers, with no gender or age differences. Occupational stress, sleep quality, and mental health showed no significant gender differences. Older workers reported bad quality of sleep, occupational stress and psychological disorders more frequently than others. Metabolic syndrome was more frequent in males than in females, and in older rather than younger workers.

**Table 1.** Prevalence of syncope, presyncope and falls, and distribution of occupational distress, low quality of sleep, impaired mental health, and metabolic syndrome according to gender and age.

|                                   | Total<br>Number (%) | Male<br>Number (%) | Female<br>Number (%) | Chi<br>square<br><i>p</i> | Younger <sup>4</sup><br>Number (%) | Older <sup>5</sup><br>Number (%) | Chi<br>square<br><i>p</i> |
|-----------------------------------|---------------------|--------------------|----------------------|---------------------------|------------------------------------|----------------------------------|---------------------------|
| Syncope                           | 103 (13.9)          | 70 (7.6)           | 83 (17.3)            | 0.000                     | 84 (16.0)                          | 19 (8.8)                         | 0.011                     |
| Presyncope                        | 200 (27.0)          | 48 (18.3)          | 152 (31.7)           | 0.000                     | 151 (28.7)                         | 49 (22.8)                        | 0.100                     |
| Fall                              | 272 (36.7)          | 92(35.1)           | 180 (37.6)           | 0.506                     | 185 (35.2)                         | 87 (40.5)                        | 0.175                     |
| Fall unknown<br>origin            | 76 (10.3)           | 22 (8.4)           | 54 (11.3)            | 0.217                     | 53 (10.1)                          | 23 (10.7)                        | 0.800                     |
| Distressed <sup>1</sup>           | 278 (38.6)          | 110 (42.5)         | 168 (36.4)           | 0.106                     | 182 (35.6)                         | 96 (45.7)                        | 0.011                     |
| Bad sleeper <sup>2</sup>          | 360 (48.6)          | 120 (45.8)         | 240 (50.1)           | 0.263                     | 234 (44.5)                         | 126 (58.6)                       | 0.000                     |
| Low mental<br>health <sup>3</sup> | 152 (20.5)          | 51 (19.5)          | 101 (21.1)           | 0.592                     | 97 (18.5)                          | 55 (25.6)                        | 0.030                     |
| Metabolic syn-<br>drome           | 91 (12.3)           | 46 (17.6)          | 45 (9.4)             | 0.001                     | 43 (8.2)                           | 48 (22.3)                        | 0.000                     |

**Notes.** <sup>1</sup>Effort reward imbalance  $ERI \geq 1$ ; <sup>2</sup>Pittsburgh Sleep Quality Index  $PSQI \geq 5$ ; <sup>3</sup>General Health Questionnaire  $GHQ12 \geq 3$ ; <sup>4</sup>age <55 years; <sup>5</sup>age  $\geq 55$  years.

Most workers reported only one syncope; 37 workers (35.9%) reported 2-4 episodes and 13 (12.6%) reported 5 or more syncope recurrences. In 18 cases, syncope had occurred in the previous two years. Only 12 workers reported syncope occurrence during working activity.

As indicated in Table 2, workers who reported syncope or presyncope were found to have a lower quality of sleep, a reduced level of mental health and greater occupational distress than the other workers. The mean scores of workers with recurrent syncope differed the most from the mean values of the group. Workers reporting falls of unknown origin had lower levels of sleep quality and mental health than the group but did not report higher occupational stress (Table 2).

**Table 2.** Comparison of mean values of stress, sleep quality and mental health in workers with or without syncope, presyncope and falls of unknown origin. (Student's t test).

|                   | Stress<br>(ERI)             | Sleep Quality<br>(PSQI)     | Mental health<br>(GHQ-12)   |
|-------------------|-----------------------------|-----------------------------|-----------------------------|
| Syncope           | 1.05±0.46 vs. 0.90±0.43 *** | 6.62±4.32 vs. 4.81±3.23 *** | 2.48±3.32 vs. 1.33±2.32 *** |
| Recurrent syncope | 1.15±0.51 vs. 0.91±0.42 *** | 7.72±4.54 vs. 4.87±3.28 *** | 3.34±3.89 vs. 1.36±2.33 *** |
| Recent syncope    | 1.17±0.47 vs. 0.92±0.43 *   | 7.56±5.01 vs. 5.00±3.39 **  | 2.67±2.97 vs. 1.46±2.49 *   |

|                       |                             |                             |                             |
|-----------------------|-----------------------------|-----------------------------|-----------------------------|
| Presyncope            | 1.06±0.46 vs. 0.88±0.41 *** | 6.88±3.97 vs. 4.39±2.98 *** | 2.40±3.21 vs. 1.16±2.10 *** |
| Fall of unknown cause | 0.97±0.46 vs. 0.92±0.43     | 6.22±4.28 vs. 4.93±3.33 **  | 2.33±3.57 vs. 1.40±2.34 **  |

Notes. \*p<0.05; \*\*p<0.01; \*\*\*p<0.001.

The levels of HDL-cholesterol triglycerides, blood glucose and blood pressure in workers with one or more T-LOC episodes did not differ from those measured in other workers (Table 3).

**Table 3.** Comparison of mean values of systolic and diastolic blood pressure, HDL-cholesterol, triglycerides, blood glucose and BMI in workers with or without T-LOC. (Student's t test).

|                       | Systolic blood pressure       | Diastolic blood pressure    | HDL-cholesterol               |
|-----------------------|-------------------------------|-----------------------------|-------------------------------|
| Syncope               | 122.35±16.40 vs. 122.28±16.29 | 79.09±13.97 vs. 79.00±11.96 | 63.86±15.48 vs. 61.96±16.22   |
| Recurrent syncope     | 122.00±16.74 vs. 122.31±16.27 | 77.77±12.26 vs. 79.11±12.24 | 65.96±15.64 vs. 61.89±16.13   |
| Presyncope            | 121.66±16.63 vs. 122.51±16.19 | 79.34±13.20 vs. 79.90±11.91 | 65.96±15.64 vs. 61.89±16.13   |
| Fall of unknown cause | 119.59±18.17 vs. 122.58±16.07 | 77.84±13.92 vs. 79.13±12.06 | 59.06±16.56 vs. 62.63±16.04   |
|                       | Triglycerides                 | Blood glucose               | BMI                           |
| Syncope               | 93.02±54.72 vs. 102.67±54.26  | 87.41±16.43 vs. 91.14±14.23 | 23.25±3.32 vs. 25.03±4.38 *** |
| Recurrent syncope     | 86.85±60.70 vs. 102.59±53.65  | 86.84±21.42 vs. 90.92±13.86 | 24.10±4.03 vs. 24.82±4.30     |
| Presyncope            | 98.26±52.51 vs. 102.41±55.09  | 88.53±14.53 vs. 91.36±14.59 | 24.55±4.57 vs. 24.85±4.17     |
| Fall of unknown cause | 114.85±50.77 vs. 99.74±54.64  | 88.97±15.39 vs. 90.79±14.52 | 25.87±5.04 vs. 24.65±4.18 *   |

Notes. \* p<0.05; \*\* p<0.01; \*\*\* p<0.001.

Workers with one or more syncopal episodes had a significantly lower than average BMI. On the other hand, workers with falls of unknown origin, had a higher than average BMI (Table 3).

Table 4 indicates the results of logistic regression in models adjusted for age and gender. Syncope, presyncope and unexplained falls occurring during working activity were associated with occupational distress, low sleep quality, low mental health and metabolic syndrome. Interestingly, syncope and presyncope were associated with an increased risk of occupational distress, sleep problems and poor mental health, whereas no association was found between syncope, presyncope or falls without apparent cause and a diagnosis of metabolic syndrome (Table 3).

**Table 4.** Association of episodes of syncope, presyncope and falls with occupational distress, poor sleep quality, impaired mental health and metabolic syndrome (logistic regression models adjusted by age and gender).

|                    | Distress<br>OR (CI95%) | Bad sleep<br>OR (CI95%) | Low mental health<br>OR (CI95%) | MetS<br>OR (CI95%) |
|--------------------|------------------------|-------------------------|---------------------------------|--------------------|
| Syncope            | 1.62 (1.05; 2.52) *    | 1.79 (1.16; 2.77) ***   | 2.43 (1.52; 3.87) ***           | 0.61 (0.27; 1.39)  |
| Recurrent syncope  | 2.11 (1.15; 3.88) *    | 2.19 (1.18; 4.04) *     | 3.88 (2.12; 7.08) ***           | 1.02 (0.40; 2.75)  |
| Recent syncope     | 1.89 (0.71; 5.04)      | 1.68 (0.63; 4.45)       | 2.55 (0.96; 6.78)               | 1.81 (0.48; 6.78)  |
| Presyncope         | 1.77 (1.25; 2.49) ***  | 2.95 (2.08; 4.18) ***   | 2.61 (1.78; 3.84) ***           | 1.21 (0.71; 2.04)  |
| Fall unknown cause | 1.00 (0.61; 1.66)      | 1.49 (0.91; 2.42)       | 1.69 (0.99; 2.87)               | 1.33 (0.67; 2.65)  |

Notes. OR: odds ratio; CI95%: confidence interval at 95%; Distress: ERI≥1; 2; Bad sleep: PSQI≥5; Low mental health: GHQ-12≥3; MetS: metabolic syndrome, three or more components (hypertension, hyperglycaemia, low HDH cholesterol, hypertriglyceridemia, obesity). \* p<0.05; \*\* p<0.01; \*\*\* p<0.001.

#### 4. Discussion

This study, which, to the best of our knowledge, is the first conducted in the workplace to assess the prevalence of syncope, presyncope or falls with no apparent cause, indicates that these phenomena are quite common and can also occur during the performance of work activities, thus potentially endangering not only the safety of the worker, but also that of colleagues or clients and the continuity of the production process.

The loss of consciousness in our study involved about one worker in seven, with a greater frequency in females and younger workers. Presyncopal symptoms were much

more common, affecting more than one in four workers, and falls for no apparent reason affected one in ten. In our sample lifetime prevalence was lower than that observed in the general population, where it is estimated to be 42%, with a higher percentage between 10 and 30 years of age, mainly of vasovagal syncope [1, 8, 41]. In occupational cohorts, prevalence rates have been reported to be as high as 35% in healthcare workers [20], 39% among medical students [42] and 41% in air force employees (with recurrent syncope occurring in 13.5%) [43]. The disparity between our data and those reported in the general population may be explained by the structure of the age groups in our sample. Syncope typically follows a tri-modal distribution in both sexes, with an increase in cases before 20, around 60 and over 80 years of age [44, 45]. In our cohort, the third modal peak was missing and the first was based on few participants. Most of the episodes had occurred in younger workers, while those reported by middle-aged workers had often occurred many years previously. The higher prevalence of syncopal episodes in women corresponded to data reported in the literature [46]. Women are younger than men at the time of their first syncope, have lower baseline systolic blood pressure according to data reported in patients with constitutional hypotension [5], experience heat as a more common trigger, have physical symptomatology more frequently (e.g., feeling warm, seizures and greater post-syncope fatigue) and are more prone to recurrent syncope [47]. These differences in terms of age and clinical presentation may explain the higher prevalence in females in this occupational cohort.

In our cohort, syncope and presyncope were significantly associated with occupational distress, low quality of sleep and poor mental health, while no association with metabolic risk factors and cardiovascular risk was observed. On the basis of these epidemiological findings, it appears that the syncopal and presyncopal episodes observed in active workers were predominantly attributable to neuro-mediated mechanisms rather than severe cardiac conditions. Our study data are in keeping with the hypothesis of central mechanisms involved in the reflex syncope reported by Mosqueda-Garcia [22] which focuses specifically on the role of the occupational environment (Figure 1). In fact, workers are a special category in which the presence of serious heart disease can lead to removal from occupational risk and early retirement. This phenomenon, known as the "healthy worker effect" causes a lower prevalence of severe heart disease in the workforce than in the general population. Nonetheless, we cannot rule out that some of the reported episodes may have been caused by heart disease. For this reason, all workers who reported syncopal episodes were advised to see their General Practitioner to obtain a definite diagnosis, in accordance with National Health plan indications.

The sudden and brief loss of consciousness that many workers reported may be due to a number of different diseases such as orthostatic hypotension (due to drugs, hypovolemia, primary or secondary autonomic failure, others), neurally-mediated syncope, cardiogenic syncope, and less frequently to other neurologic disorders such as epilepsy, psychogenic syncope, vertebrobasilar transient ischemic attacks), metabolic disorders and intoxication [8, 48]. According to the literature, vasovagal is the most common form of syncope [49]. It accounts for 60-80% of cases of syncope, and typically occurs in young adults [4, 50]. The absence of structural heart disease in our sample and the higher prevalence in younger rather than older workers suggest that most of the reported episodes may be attributed to this benign form. Although vasovagal syndrome is generally a benign condition from a clinical point of view, it may have a negative impact on work. Indeed, it may create a danger for workers and third parties [11] if it occurs during highly hazardous tasks and may result in a significantly increased economic burden. Recurrent cases of syncope may reduce the quality of life and promote occupational injury [11]. Furthermore, they are associated with an increased risk of death and major adverse cardiovascular events [51, 52]. Atypical vasovagal syncope, which is commoner in older adults, and non-neurogenic syncope can often be erroneously misdiagnosed as falls. We agree with Kenny [49] that a more standardized approach should be adopted in the diagnosis and management of workers presenting with syncope or unexplained falls.

In this study we observed an association between syncope occurrence and occupational stress. A previous study reported that emotional vasovagal syncope might be associated with distress [53]. In addition, a longitudinal study showed that psychosocial impairment reliably predicted non-response to treatment of syncope [54]. This finding supports the hypothesis that psychosocial occupational stress factors may have an important role in syncope occurrence, particularly in hypersusceptible individuals.

The association of syncope with low sleep quality observed in our study, was not completely unexpected. Indeed, daytime sleepiness due to low quality of night sleep has been associated with orthostatic hypotension that may promote syncope, presyncope and falls [55]. Sporadic observations in the literature suggest a link between sleep apnoea syndrome and vasovagal syncope [56]. Interestingly, sleep quality is a mediator between occupational stress and its effects on metabolic pathologies [57] and mental health [58]. Vasovagal syncope has also been previously associated with mental health problems [59]. In a longitudinal study, syncope patients exhibited high levels of psychological distress and mood/anxiety disorders [60]. Psychiatric disorders are common in patients with tilt-induced vasovagal syncope and seem to predict the risk of recurrence [59].

A significant share of the workers in our sample had recurrent episodes of syncope. There is considerable evidence that like other chronic diseases, syncope recurrence affects the quality of life [61] by impacting negatively on numerous daily activities such as driving, working, attending school, and by impairing mental health with episodes of somatization, depression and anxiety [62-64].

This study has some limitations. The use of a convenience sample leads to a very cautious application of the results to other occupational situations. However, our research showed that searching for T-LOC events among workers is useful, achievable with limited use of resources and does not burden overly the health and safety services. For this reason, other future studies might reinforce the evidence. Another limitation is the cross-sectional nature of the study, which does not allow to infer causality. Furthermore, the episodes of T-LOC are self-reported, and this allows to exclude neither underreporting nor overreporting.

## 5. Conclusions

The association of symptoms with excessive occupational stress, low quality of sleep and mental balance disturbances indicates the need to consider health promotion intervention in the workplace. A prevention program should aim at improving sleep quality, reducing stressors and increasing worker resilience. Psychological support should be provided, at least for workers with more evident mental health problems.

**Author Contributions:** Conceptualization, N.M.; methodology, N.M. and F.B.; formal analysis, N.M. and R.R.D.P.; investigation, N.M., G.A., S.G., A.I., I.M., A.C., M.Gab., M.Gas., M.L., C.M.; writing, original draft preparation, N.M.; writing, review and editing, N.M., and F.B. All authors have read and agreed to the published version of the manuscript."

**Funding:** This research received no external funding.

**Institutional Review Board Statement:** The study was conducted according to the guidelines of the Declaration of Helsinki and approved by the Institutional Review Board of the Department of Woman, Child & Public Health of the Università Cattolica del Sacro Cuore.

**Informed Consent Statement:** Informed consent was obtained from all subjects involved in the study.

**Data Availability Statement:** The data presented in this study are freely available on Zenodo repository.

**Acknowledgments:** We thanks Ms. Elisabeth Ann Wright who revised the English language.

**Conflicts of Interest:** The authors declare no conflict of interest.

## References

- Shen, W.K.; Sheldon, R.S.; Benditt, D.G.; Cohen, M.I.; Forman, D.E.; Goldberger, Z.D.; Grubb, B.P.; Hamdan, M.H.; Krahn, A.D.; Link, M.S.; et al. 2017 ACC/AHA/HRS Guideline for the Evaluation and Management of Patients With Syncope: A Report of the American College of Cardiology/American Heart Association Task Force on Clinical Practice Guidelines and the Heart Rhythm Society. *Circulation* **2017**, *136*(5), e60-e122. doi: 10.1161/CIR.0000000000000499. Erratum in: *Circulation* **2017**, *136*(16), e271-e272.
- Brignole, M.; Moya, A.; de Lange, F.J.; Deharo, J.C.; Elliott, P.M.; Fanciulli, A.; Fedorowski, A.; Furlan, R.; Kenny, R.A.; Martín, A.; et al. 2018 ESC Guidelines for the diagnosis and management of syncope. *Eur Heart J* **2018**, *39*(21), 1883-1948. doi: 10.1093/eurheartj/ehy037.
  - Mosqueda-Garcia, R.; Furlan, R.; Tank, J.; Fernandez-Violante, R. The elusive pathophysiology of neurally mediated syncope. *Circulation* **2000**, *102*(23), 2898-906. doi: 10.1161/01.cir.102.23.2898.
  - Barbic, F.; Heusser, K.; Minonzio, M.; Shiffer, D.; Cairo, B.; Tank, J.; Jordan, J.; Diedrich, A.; Gauger P.; Zamuner RA.; et al. Effects of Prolonged Head-Down Bed Rest on Cardiac and Vascular Baroreceptor Modulation and Orthostatic Tolerance in Healthy Individuals. *Front Physiol* **2019**, *10*, 1061. doi: 10.3389/fphys.2019.01061.
  - Jacob, G.; Barbic, F.; Glago, M.; Dipaola, F.; Porta, A.; Furlan, R. Cardiovascular autonomic profile in women with constitutional hypotension. *J Hypertens* **2018**, *36*(10), 2068-2076. doi: 10.1097/HJH.0000000000001790.
  - Goldberger, Z.D.; Petek, B.J.; Brignole, M.; Shen, W.K.; Sheldon, R.S.; Solbiati, M.; Deharo, J.C.; Moya, A.; Hamdan, M.H. ACC/AHA/HRS Versus ESC Guidelines for the Diagnosis and Management of Syncope: JACC Guideline Comparison. *J Am Coll Cardiol* **2019**, *74*(19), 2410-2423. doi: 10.1016/j.jacc.2019.09.012.
  - Alciati, A.; Shiffer, D.; Dipaola, F.; Barbic, F.; Furlan, R. Psychogenic Pseudosyncope: Clinical Features; Diagnosis and Management. *J Atr Fibrillation* **2020**, *13*(1), 2399. doi: 10.4022/jafib.2399.
  - Task Force for the Diagnosis and Management of Syncope; European Society of Cardiology (ESC); European Heart Rhythm Association (EHRA); Heart Failure Association (HFA); Heart Rhythm Society (HRS); Moya, A.; Sutton, R.; Ammirati, F.; Blanc, J.J.; Brignole, M.; et al. Guidelines for the diagnosis and management of syncope (version 2009). *Eur Heart J* **2009**, *30*(21), 2631-71. doi: 10.1093/eurheartj/ehp298.
  - Bhangu, J.; Hall, P.; Devaney, N.; Bennett, K.; Carroll, L.; Kenny, R.A.; McMahon, C.G. The prevalence of unexplained falls and syncope in older adults presenting to an Irish urban emergency department. *Eur J Emerg Med* **2019**, *26*(2), 100-104. doi: 10.1097/MEJ.0000000000000548.
  - Sun, B.C.; Costantino, G.; Barbic, F.; Bossi, I.; Casazza, G.; Dipaola, F.; McDermott, D.; Quinn, J.; Reed, M.; Sheldon, R.S.; et al. Priorities for emergency department syncope research. *Ann Emerg Med* **2014**, *64*(6), 649-55.e2. doi: 10.1016/j.annemerg-med.2014.04.014.
  - Barbic, F.; Casazza, G.; Zamuner, A.R.; Costantino, G.; Orlandi, M.; Dipaola, F.; Capitano, C.; Achenza, S.; Sheldon, R.; Furlan, R. Driving and working with syncope. *Auton Neurosci* **2014**, *184*, 46-52. doi: 10.1016/j.autneu.2014.05.006.
  - Barbic, F.; Dipaola, F.; Casazza, G.; Borella, M.; Minonzio, M.; Solbiati, M.; Raj, S.R.; Sheldon, R.; Quinn, J.; Costantino, G.; et al. Syncope in a Working-Age Population: Recurrence Risk and Related Risk Factors. *J Clin Med* **2019**, *8*(2), 150. doi: 10.3390/jcm8020150.
  - Numé, A.K.; Kragholm, K.; Carlson, N.; Kristensen, S.L.; Bøggild, H.; Hlatky, M.A.; Torp-Pedersen, C.; Gislason, G.; Ruwald, M.H. Syncope and Its Impact on Occupational Accidents and Employment: A Danish Nationwide Retrospective Cohort Study. *Circ Cardiovasc Qual Outcomes* **2017**, *10*(4), e003202. doi: 10.1161/CIRCOUTCOMES.116.003202.
  - Armed Forces Health Surveillance Center (AFHSC). Syncope, active and reserve components, U.S. Armed Forces, 1998-2012. *MSMR* **2013**, *20*(11), 5-9.
  - Rudnicki, J.; Zyśko, D.; Gajek, J.; Kuliczowski, W.; Rosińczuk-Tonderys, J.; Zielińska, D.; Terpiłowski, Ł.; Agrawal, A.K. The risk for syncope and presyncope during surgery in surgeons and nurses. *Pacing Clin Electrophysiol* **2011**, *34*(11), 1486-91. doi: 10.1111/j.1540-8159.2011.03169.x.
  - Soteriades, E.S.; Evans, J.C.; Larson, M.G.; Chen, M.H.; Chen, L.; Benjamin, E.J.; Levy, D. Incidence and prognosis of syncope. *N Engl J Med* **2002**, *347*(12), 878-85. doi: 10.1056/NEJMoa012407.
  - Ganzeboom, K.S.; Mairuhu, G.; Reitsma, J.B.; Linzer, M.; Wieling, W.; van Dijk, N. Lifetime cumulative incidence of syncope in the general population: a study of 549 Dutch subjects aged 35-60 years. *J Cardiovasc Electrophysiol* **2006**, *17*(11), 1172-6. doi: 10.1111/j.1540-8167.2006.00595.x.
  - Parsons, I.T.; Cox, A.T.; Mollan, I.A.; Boos, C.J. Managing the military patient with syncope. *J R Army Med Corps* **2015**, *161*(3), 180-6. doi: 10.1136/jramc-2015-000493.
  - Barbic, F.; Borella, M.; Perego, F.; Dipaola, F.; Costantino, G.; Galli, A.; Mantovani, C.; Seghizzi, P.; Malliani, A.; Furlan, R. La sincope in età lavorativa. Studio multicentrico prospettico STePS [Syncope and work. STePS study (Short Term Prognosis of Syncope)]. *G Ital Med Lav Ergon* **2005**, *27*(3), 272-4.
  - Gaggioli, G.; Laffi, M.; Montemanni, M.; Mocini, A.; Rubartelli, P.; Brignole, M. Risk of syncope during work. *Europace*. **2014**, *16*(2), 289-92. doi: 10.1093/europace/eut247.
  - Furlan, R.; Alboni, P.; Mosqueda-Garcia, R. Pathophysiology of vasovagal syncope: Conclusive remarks. In *Vasovagal Syncope*; 1st ed.; Alboni, P.; Furlan, R. Editors Springer International Publishing, Switzerland, 2015; pp. 95-102. doi: 10.1007/978-3-319-09102-0\_9.
  - Mosqueda-Garcia R. Role of Autonomic Nervous System in Vasovagal Syncope. *Vasovagal Syncope*, 1st ed.; Alboni, P.; Furlan, R. Editors; Springer International Publishing: Switzerland, 2015; pp 53-65. doi: 10.1007/978-3-319-09102-0\_5.
  - Solbiati, M.; Casazza, G.; Dipaola, F.; Rusconi, AM.; Cernuschi, G.; Barbic, F.; Montano, N.; Sheldon, R.S.; Furlan, R.; Costantino, G. Syncope recurrence and mortality: a systematic review. *Europace* **2015**, *17*(2), 300-8. doi: 10.1093/europace/euu327.

23. Magnavita, N. Two tools for health surveillance of job stress: the Karasek Job Content Questionnaire and the Siegrist Effort Reward Imbalance Questionnaire. *G Ital Med Lav Ergon* **2007**, *29*(3), 667–670.
24. Magnavita, N.; Garbarino, S.; Siegrist, J. The use of parsimonious questionnaires in occupational health surveillance. Psychometric properties of the short Italian version of the Effort/Reward Imbalance questionnaire. *TSWJ Sci World J* **2012**, *2012*, 372852. doi: 10.1100/2012/372852.
25. Siegrist, J. Adverse health effects of high-effort/low-reward conditions. *J Occup Health Psychol* **1996**, *1*(1), 27–41.
26. Siegrist, J.; Wege, N.; Puhlhofer, F.; Wahrendorf, M. A short generic measure of work stress in the era of globalization: effort-reward imbalance. *Int Arch Occup Environ Health* **2009**, *82*(8), 1005-13.
27. Curcio, G.; Tempesta, D.; Scarlata, S.; Marzano, C.; Moroni, F.; Rossini, P.M.; Ferrara, M.; De Gennaro, L. Validity of the Italian version of the Pittsburgh Sleep Quality Index (PSQI). *Neurol Sci* **2013**, *34*(4), 511-9. doi: 10.1007/s10072-012-1085-y.
28. Buysse, D.J.; Reynolds, C.F.; Monk, T.H.; Berman, S.R.; Kupfer, D.J. The Pittsburgh Sleep Quality Index (PSQI): A new instrument for psychiatric research and practice. *Psychiatry Res* **1989**, *28*(2), 193-213.
29. Magnavita, N. *Lavoro umano. Il benessere nei luoghi di lavoro*; EDUCatt: Milano, Italy, 2009; ISBN 978-88-8311-722-0.
30. Piccinelli, M.; Bisoffi, G.; Bon, M.G.; Cunico, L.; Tansella, M. Validity and test-retest reliability of the Italian version of the 12-item GHQ in general practice: a comparison between three scoring methods. *Compr Psychiatry* **1993**, *34*, 198-205.
31. Goldberg, D. *The detection of psychiatric illness by questionnaire*; Oxford University Press: London, United Kingdom, 1972.
32. Goldberg, D.; Hillier, V.F. A scaled version of the General Health Questionnaire. *Psychol Med* **1979**, *9*(1), 139-45.
33. Banks, M.H.; Clegg, C.W.; Jackson, P.R.; Kemp, N.J.; Stafford, E.M.; Wall, T.D. The use of the General Health Questionnaire as an indicator of mental health in occupational studies. *J Occup Psychol* **1980**, *53*, 187-194.
34. Banks, M.H. Validation of the General Health Questionnaire in a young community sample. *Psychol Med* **1983**, *13*, 349-353.
35. The IDF consensus worldwide definition of the metabolic syndrome. Available online: <https://www.idf.org/e-library/consensus-statements/60-idfconsensus-worldwide-definition-of-the-metabolic-syndrome.html> (accessed on 13 September 2021).
36. Expert panel on detection, evaluation, and treatment of high blood cholesterol in adults. Executive summary of the third report of the National Cholesterol Education Program (NCEP) expert panel on detection, evaluation, and treatment of high blood cholesterol in adults (adult treatment panel III). *JAMA* **2001**, *285*(19), 2486-97. doi: 10.1001/jama.285.19.2486.
37. Einhorn, D.; Reaven, G.M.; Cobin, R.H.; Ford, E.; Ganda, O.P.; Handelsman, Y.; Hellman, R.; Jellinger, P.S.; Kendall, D.; Krauss, R.M.; et al. American College of Endocrinology position statement on the insulin resistance syndrome. *Endocr Pract* **2003**, *9*(3), 237-52.
38. Kassi, E.; Pervanidou, P.; Kaltsas, G.; Chrousos, G. Metabolic syndrome: definitions and controversies. *BMC Med* **2011**, *9*, 48. doi:10.1186/1741-7015-9-48.
39. Poscia, A.; Moscato, U.; La Milia, D.I.; Milovanovic, S.; Stojanovic, J.; Borghini, A.; Collamati, A.; Ricciardi, W.; Magnavita, N. Workplace Health Promotion for Older Workers: a Systematic Literature Review. *BMC Health Serv Res* **2016**, *16* Suppl 5, 329. doi: 10.1186/s12913-016-1518-z.
40. da Silva, R.M. Syncope: epidemiology, etiology, and prognosis. *Front Physiol* **2014**, *5*, 471. doi: 10.3389/fphys.2014.00471.
41. Serletis, A.; Rose, S.; Sheldon, A.G.; Sheldon, R.S. Vasovagal syncope in medical students and their first-degree relatives. *Eur Heart J* **2006**, *27*(16), 1965-70. doi: 10.1093/eurheartj/ehl147.
42. Lamb, L.E. Incidence of loss of consciousness in 1980 Air Force personnel. *Aerosp Med* **1960**, *31*, 973–88.
43. Ganzeboom, K.S.; Colman, N.; Reitsma, J.B.; Shen, W.K.; Wieling, W. Prevalence and triggers of syncope in medical students. *Am J Cardiol.* **2003**;91(8):1006-8, A8. doi: 10.1016/s0002-9149(03)00127-9.
44. Ruwald, M.H.; Hansen, M.L.; Lamberts, M.; Hansen, C.M.; Højgaard, M.V.; Køber, L.; Torp-Pedersen, C.; Hansen, J.; Gislason, G.H. The relation between age, sex, comorbidity, and pharmacotherapy and the risk of syncope: a Danish nationwide study. *Europace* **2012**, *14*(10), 1506-14. doi: 10.1093/europace/eus154.
45. Chen, L.Y.; Shen, W.K.; Mahoney, D.W.; Jacobsen, S.J.; Rodeheffer, R.J. Prevalence of syncope in a population aged more than 45 years. *Am J Med* **2006**, *119*(12), 1088.e1-7. doi: 10.1016/j.amjmed.2006.01.029.
46. Deveau, A.P.; Sheldon, R.; Maxey, C.; Ritchie, D.; Doucette, S.; Parkash, R. Sex Differences in Vasovagal Syncope: A Post Hoc Analysis of the Prevention of Syncope Trials (POST) I and II. *Can J Cardiol* **2020**, *36*(1), 79-83. doi: 10.1016/j.cjca.2019.10.008.
47. Bassetti, C.L. Transient loss of consciousness and syncope. *Handb Clin Neurol* **2014**, *119*, 169-91. doi: 10.1016/B978-0-7020-4086-3.00013-8.
48. Kenny, R.A.; McNicholas, T. The management of vasovagal syncope. *QJM* **2016**, *109*(12), 767-773. doi: 10.1093/qjmed/hcw089.
49. Li, H.X.; Gao, L.; Yuan, Y. Advance in the understanding of vasovagal syncope in children and adolescents. *World J Pediatr* **2021**, *17*(1), 58-62. doi: 10.1007/s12519-020-00367-z.
50. Zimmermann, T.; du Fay de Lavallaz, J.; Nestelberger, T.; Gualandro, D.M.; Strebel, I.; Badertscher, P.; Lopez-Ayala, P.; Widmer, V.; Freese, M.; Miró, Ò.; et al. Incidence, characteristics, determinants, and prognostic impact of recurrent syncope. *Europace* **2020**; *22*(12), 1885-1895. doi: 10.1093/europace/euaa227.
51. Jorge, J.G.; Pournazari, P.; Raj, S.R.; Maxey, C.; Sheldon, R.S. Frequency of injuries associated with syncope in the prevention of syncope trials. *Europace* **2020**, *22*(12), 1896-1903. doi: 10.1093/europace/euaa246.
52. Zysko, D.; Melander, O.; Fedorowski, A. Vasovagal syncope related to emotional stress predicts coronary events in later life. *Pacing Clin Electrophysiol* **2013**, *36*(8), 1000-6. doi: 10.1111/pace.12138.
53. Flint, B.; Baker, C.; Freeston, M.; Newton, J.L. Level of psychosocial impairment predicts early response to treatment in vasovagal syncope. *Europace* **2009**, *11*(2), 231-6. doi: 10.1093/europace/eun332.

54. Lewis, N.C.; Jones, H.; Ainslie, P.N.; Thompson, A.; Marrin, K.; Atkinson, G. Influence of nocturnal and daytime sleep on initial orthostatic hypotension. *Eur J Appl Physiol* **2015**, *115*(2), 269-76. doi: 10.1007/s00421-014-3010-y.
55. Puel, V.; Godard, I.; Papaioannou, G.; Gosse, P.; Pepin, J.L.; Thoin, F.; Deharo, J.C.; Roche, F.; Zarqane, N.; Gagnadoux, F.; et al. Management of sleep apnoea syndrome (SAS) in patients with vasovagal syncope (VVS): a protocol for the VVS-SAS cohort study. *BMJ Open* **2020**, *10*(9), e038791. doi: 10.1136/bmjopen-2020-038791.
56. Garbarino, S.; Magnavita, N. Sleep problems are a strong predictor of stress-related metabolic changes in police officers. A prospective study. *PLoS One* **2019**, *14*(10), e0224259. doi: 10.1371/journal.pone.0224259.
57. Magnavita, N.; Tripepi, G.; Di Prinzio, R.R. Symptoms in Health Care Workers during the COVID-19 Epidemic. A Cross-Sectional Survey. *Int J Environ Res Public Health* **2020**, *17*(14), 5218. doi: 10.3390/ijerph17145218.
58. Giada, F.; Silvestri, I.; Rossillo, A.; Nicotera, P.G.; Manzillo, G.F.; Raviele, A. Psychiatric profile quality of life and risk of syncope recurrence in patients with tilt-induced vasovagal syncope. *Europace* **2005**, *7*(5), 465-71. doi: 10.1016/j.eupc.2005.05.008.
59. D'Antono, B.; Dupuis, G.; St-Jean, K.; Lévesque, K.; Nadeau, R.; Guerra, P.; Thibault, B.; Kus, T. Prospective evaluation of psychological distress and psychiatric morbidity in recurrent vasovagal and unexplained syncope. *J Psychosom Res* **2009**, *67*(3), 213-22. doi: 10.1016/j.jpsychores.2009.03.012.
60. Picavet, H.S.; Hoeymans, N. Health related quality of life in multiple musculoskeletal diseases: SF-36 and EQ-5D in the DMC3 study. *Ann Rheum Dis* **2004**, *63*(6), 723-9. doi: 10.1136/ard.2003.010769.
61. Anderson, J.B.; Czosek, R.J.; Knilans, T.K.; Marino, B.S. The effect of paediatric syncope on health-related quality of life. *Cardiol Young* **2012**, *22*(5), 583-8. doi: 10.1017/S1047951112000133.
62. Rose, M.S.; Koshman, M.L.; Ritchie, D.; Sheldon, R. The development and preliminary validation of a scale measuring the impact of syncope on quality of life. *Europace* **2009**, *11*(10), 1369-74. doi: 10.1093/europace/eup106.
63. van Dijk, N.; Sprangers, M.A.; Colman, N.; Boer, K.R.; Wieling, W.; Linzer, M. Clinical factors associated with quality of life in patients with transient loss of consciousness. *J Cardiovasc Electrophysiol* **2006**, *17*(9), 998-1003. doi: 10.1111/j.1540-8167.2006.00533.x.