

Prevalence of Sleep Disordered Breathing Symptoms and Their Relation with Concomitant Diseases in Afyon, Turkey: A Population Based Study

Afyon İlinde Uykuda Solunum Bozukluğu Semptom Prevalansı ve Bu Semptomların Eşlik Eden Hastalıklarla İlişkisi: Toplum Temelli Çalışma

Mehmet ÜNLÜ, MD,^a
Murat SEZER, MD,^a
Fatma FİDAN, MD,^a
Abdullah AYÇİÇEK, MD,^b
Dilek TOPRAK, MD,^c
Nurhan DOĞAN, MD,^d
Ziya KARA, MD^a

Departments of ^aPulmonary Medicine, ^bEar, Nose and Throat, ^cFamily Medicine, ^dBioStatistics, Afyonkarahisar Kocatepe University, Faculty of Medicine, Afyonkarahisar

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Yazışma Adresil/Correspondence:
Murat SEZER, MD
Afyonkarahisar Kocatepe University
Faculty of Medicine,
Department of Pulmonary Medicine,
Afyonkarahisar,
TÜRKİYE/TURKEY
drrmuratsezer@yahoo.com

ABSTRACT Objective: To estimate the prevalence of symptoms of sleep-disordered breathing (SDB) and to evaluate the relationship between these symptoms and concomitant systemic disorders in Turkish adults. **Material and Methods:** A sample group representing whole Afyon city filled, a questionnaire consisting of questions about demographic features, SDB symptoms and concomitant diseases. Body mass indices of the subjects were calculated. **Results:** There was no significant difference for snoring between males and females, whereas witnessed apnea and excessive daytime sleepiness were significantly higher in females. Obesity was more prevalent among females. SDB symptoms were more common among obese subjects. Diabetes (DM), hypertension (HT), hyperlipidemia and congestive heart failure (CHF) were significantly more prevalent among snorers. There was a 1.7 folds increase in HT risk and a 1.4 folds increase in DM risk in snorers. We also found a 1.4 folds increased risk for hyperlipidemia and 2.1 folds increased risk for CHF in our snorer population. Snoring and witnessed apnea were found as risk factors for hypertension independently from smoking status, BMI and age. **Conclusion:** The frequency of SDB symptoms were high in Afyon city, with a greater than expected rate for females. Snoring and witnessed apnea were found as independent risk factors for hypertension.

Key Words: Diabetes mellitus; hypertension; prevalence; sleep apnea syndromes; snoring

ÖZET Amaç: Türk erişkinlerde uykuda solunum bozukluğu semptom prevalansını saptamak ve bu semptomlar ile eşlik eden sistemik hastalıklar arasındaki ilişkiyi incelemek. **Gereç ve Yöntemler:** Tüm Afyon ilini temsil edecek şekilde seçilmiş örneklem grubundaki kişilere demografik özellikleri, uykuda solunum bozukluğu semptomları ve eşlik eden hastalıkların sorgulandığı bir anket dolduruldu. Katılımcıların beden kitle indeksleri hesaplandı. **Bulgular:** Erkek ve kadınlar arasında horlama sıklığı açısından istatistiksel olarak anlamlı fark saptanmazken, tanıklı apne ve gündüz aşırı uyku sıklığı kadınlarda daha sık olarak bulundu. Obezite kadınlarda daha sık idi. Obez hastalarda uykuda solunum bozukluğu semptomları daha yaygın idi. Diabet (DM), hipertansiyon (HT), hiperlipidemi ve konjestif kalp yetmezliği (KKY) horlaması olanlarda anlamlı olarak daha sık saptandı. Horlaması olanlarda HT riski 1.7 kat, DM riski 1.4 kat artmış olarak bulundu. Yine horlaması olan popülasyonda hiperlipidemi riski 1.4 kat ve KKY riski 2.1 kat artmış olarak bulundu. Horlama ve tanıklı apne, hipertansiyon için; sigara içme durumu, beden kitle indeksi ve yaştan bağımsız risk faktörleri olarak tespit edildi. **Sonuç:** Uykuda solunum bozukluğu semptomları sıklığı Afyon ilinde yüksektir ve bu sıklık kadınlarda beklenenden de yüksek olarak tespit edilmiştir. Horlama ve tanıklı apne hipertansiyon için bağımsız risk faktörleri olarak tespit edilmiştir.

Anahtar Kelimeler: Diabetes mellitus; hipertansiyon; prevalans; uykuda solunum bozukluğu; horlama

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Sleep disordered breathing (SDB) is a common syndrome.^{1,2} It is associated with various medical problems³⁻⁵ that have impact on morbidity and mortality. It also causes an additional burden of the public health

service.⁶⁻⁸ There have been many epidemiologic studies to establish the prevalence of SDB and obstructive sleep apnea syndrome (OSAS) in Western countries,^{1,2,9} with limited data having been published in Asian countries.¹⁰⁻¹² Previous studies reported that the prevalence of SDB ranged from 3 to 28% in Western countries, depending on the definition of SDB and the methodology of the studies. Recently, among Asians, the prevalence of SDB was estimated to be 8.8% in Chinese office men¹¹ and 19.5% in normal Indian men.¹² Obstructive sleep apnea has long been observed in clinical practice to be more common in men,^{13,14} although it has been documented in population studies^{1,9,15} that the male dominance of sleep-disordered breathing (SDB) may not be as high, suggesting an underrepresentation and/or underrecognition of this problem in women in the clinic population. SDB may also have different prevalence and risk factors in different communities and ethnic groups. The Wisconsin sleep cohort study observed a three times higher prevalence of sleep disordered breathing (SDB) in men (24%) than in women (9%).¹ This gender difference in the prevalence of SDB has not been adequately explained but suggests that the risk factors and mechanisms for OSA may differ between men and women. It is commonly thought that hormonal factors may be one of the reasons accounting for these differences.

Hypertension and type 2 diabetes are two components of the metabolic syndrome. Apart from well-known risk factors, such as obesity, physical inactivity and excessive alcohol intake, there are also data indicating that both disorders are related to the OSAS.^{3,16-18} The underlying pathophysiologic mechanisms are not fully understood, but it can be speculated that hypoxemia, hypercapnia and/or arousal from sleep, followed by the chronic activation of the sympathetic nervous system¹⁹ negatively influence the metabolic and cardiovascular system. Snoring is one of the cardinal symptoms of OSAS. Several studies have found that snoring is positively associated with both hypertension²⁰⁻²² and diabetes.^{23,24} It seems reasonable that snoring increases the risk of hypertension and diabetes through OSAS.²⁵

The purpose of this study was to estimate the prevalence of symptoms related with sleep disordered breathing and to evaluate the relationship between snoring and systemic disorders in Turkish adults of 19-90 years age.

MATERIAL AND METHODS

The study was conducted between November 2005 and February 2006, in Afyonkarahisar, a city located in middle Anatolia in Turkey, 350 km from the capital Ankara and has a population of approximately 817000. The study was approved by the Afyon Kocatepe University Faculty of Medicine Clinical Research Ethics Committee and written, informed consent was obtained from all participants.

A total of 7000 km. roadway driven for the research by a team of 15 physicians, one nurse and a driver. A total of 2035 people, from 75 different screening regions (18 urban, 57 villages) of our city were detected with stratified sampling method, according to the population records of the year 2000, which represent the population of the area appropriately. Regions and subjects were selected regarding to represent minimum %80 of population of that village. In every research point, minimum 20 people were included to the study. The records of the regional health institutions were used in order to determine the subjects. The study group was selected randomly from the "Family Cards" of the primary health centers, regarding the gender and ages. Only one person was selected from every house. According to population distribution of year 2000, we determined the minimum number of people as 1990 (when $d=0.02$, 5% error and 95% confidence interval) and at the end of the study we reached a number of 2035 people.

The subjects were informed about the study by telephone interviews one night before, their approvals were obtained and their transport to the health institutions, where the study would be conducted, was provided. The data were collected by a questionnaire in which face to face survey method was performed by the physicians. For questions regarding symptoms related to sleep disturbances, the subjects were asked for presence of snoring, witnessed apnea and excessive day-time

sleepiness. Example questions: “Have your bedpartner or spouse ever told you that you were snoring in your sleep?”, “Have your bedpartner or spouse ever told you that you stopped breathing for some time during your sleep?”.

Body mass index was calculated according to the formula: weight/square of height. Participants with a BMI ≥ 30 were classified as obese, and those with a BMI < 30 were classified as non-obese. Participants were asked for physician diagnosed diabetes mellitus (DM), hypertension (HT), hyperlipidemia, congestive heart failure (CHF), cerebrovascular accidents (CVA) and erectile dysfunction.

STATISTICAL ANALYSIS

Statistical analyses were performed using Statistical Package for the Social Sciences (SPSS) version 12.0. The results were presented as the means \pm SD. The Chi-square test was used to compare the differences between proportions. When the comparison involved continuous variables, the t-test was used. Multiple logistic regression analysis was performed for simultaneous evaluations of more than two variables and the results were expressed as odds ratios (OR) with 95% confidence intervals (CI). To analyze the independent influence of snoring on the somatic diseases, multivariate models were used, with adjustment for age, BMI, alcohol addiction and smoking status. Age was categorized into intervals (18-24, 25-44, 45-64, 65-74, >75) in the multiple analysis, while BMI was categorized into four intervals (<25: normal, 25.0-29.9: overweight, 30.0-39.9: obese, >40: morbid obese).

RESULTS

Of the 2035 participants, 58.7% were females and 41.3% were males. Demographic features and major symptoms related to SDB according to gender were shown in Table 1. There was no significant difference between males and females for mean age. BMI was significantly higher in females than in males ($p= 0.000$). Men were smoking significantly more than women ($p= 0.000$).

Evaluation of major symptoms related to SDB revealed that there was no significant difference for snoring between males and females ($p= 0.541$), whereas witnessed apnea and excessive daytime sleepiness were significantly higher in females ($p= 0.000$). Mean age of obese subjects was significantly higher than non-obese ones ($p= 0.000$) and obesity was more prevalent among females ($p= 0.000$). Snoring, witnessed apnea and excessive daytime sleepiness were also more frequent among obese subjects (Table 2). Mean age of snorers was 49.7 ± 11.3 years and that of non-snorers was 46.9 ± 13.9 years. The difference was statistically significant

TABLE 2: Demographic features and major SDB symptoms with regard to obesity.

	Non-obese BMI<30 (n=1390)	Obese BMI ≥ 30 (n=645)	P value
Age, years (Mean \pm SD)	46.8 \pm 14.0	49.7 \pm 10.4	<0.001
Female/Male (%)	50.7/49.3	72.3/27.7	<0.001
Snoring (%)	27.1	48.8	<0.001
Witnessed apnea (%)	9.7	14.2	0.003
Excessive daytime sleepiness (%)	25.4	30.4	0.021

TABLE 1: Demographic features and major SDB symptoms with regard to gender.

	Female (n=1194)	Male (n=841)	Total (2035)	P value
Age, years, (Mean \pm SD)	47.9 \pm 12.8	47.8 \pm 13.6	47.9 \pm 13.1	0.881
BMI, kg/m ² , (Mean \pm SD)	28.9 \pm 6.1	26.5 \pm 4.5	27.9 \pm 5.6	<0.001
Smoking status				
Never (%)	91.3	25.6	64.0	
Former (%)	1.4	25.3	11.3	<0.001
Current (%)	7.2	49.1	24.6	
Alcohol consumption (%)	5.2	7.3	5.9	0.058
Snoring (%)	33.4	34.7	34.0	0.541
Witnessed apnea (%)	13.7	7.8	11.2	<0.001
Excessive daytime sleepiness (%)	30.8	21.9	27.1	<0.001

TABLE 3: Prevalence of concomitant diseases of snorers and non-snorers.

	Snorers (n= 691)	Non-snorers (n= 1344)	p value
Diabetes mellitus (%)	15.5	11.7	0.016
Hypertension (%)	31.1	20.7	<0.001
Hyperlipidemia (%)	41.0	33.3	0.001
Congestive heart failure (%)	2.9	1.4	0.021
Cerebrovascular event (%)	1.7	1.2	0.317
Erectile dysfunction (%)	6.1	4.6	0.155

($p < 0.001$). Of the snorers 57.7% were females and 42.3% were males. There was no statistically significant difference between snorers and non-snorers

according to gender ($p = 0.541$). Prevalence of concomitant diseases of snorers and non-snorers were shown in Table 3. DM (1.4 times, $p = 0.016$), HT (1.7 times, $p < 0.001$), hyperlipidemia (1.4 times, $p = 0.001$) and CHF (2.1 times, $p = 0.024$) were significantly more prevalent among snorers. CVA and erectile dysfunction were also more prevalent among snorers but the difference was not statistically significant. Risk factors for hypertension and diabetes in the multiple logistic regression analysis is shown in Table 4.

Comparison of major symptoms related to SDB according to age groups revealed that snoring was most prevalent in 45-64 years age group, wit-

TABLE 4: Risk factors for hypertension and diabetes in the multiple logistic regression analysis.

	OR (95% CI) for Hypertension		OR (95% CI) for Diabetes	
	Univariate analysis	Multivariate analysis	Univariate analysis	Multivariate analysis
Age, years				
18-24	1	1	1	1
25-44	5.12 (1.27-20.70) ($p=0.022$) (B:1.634)	2.90 (0.69-12.19) (B:1.063)*	5.27 (0.73-38.24) ($p=0.100$) (B:1.663)	3.93 (0.53-28.89) (B:1.368)
45-64	17.96 (4.51-71.56) ($p=0.000$) (B:2.888)	8.54 (2.05-35.52) (B:2.145)†	18.62 (2.62-132.53) ($p=0.003$) (B:2.924)	12.53 (1.73-90.78) (B:2.528)†
65-74	41.88 (10.28-170.66) ($p=0.000$) (B:3.735)	22.58 (5.31-96.03) (B:3.117)†	25.45 (3.50-185.28) ($p=0.001$) (B:3.237)	17.92 (2.42-132.76) (B:2.886)†
>75	48.37 (10.84-215.77) ($p=0.000$) (B:3.879)	27.35 (5.72-130.72) (B:3.309)†	13.48 (1.59-114.13) ($p=0.017$) (B:2.601)	10.96 (1.26-94.91) (B:2.394)†
BMI, kg/m²				
<25.0	1	1	1	1
25.0-29.9	1.43 (1.07-1.90) ($p=0.015$) (B:0.356)	1.18 (0.85-1.62) (B:0.161)	1.59 (1.22-2.26) ($p=0.009$) (B:0.465)	1.31 (0.90-1.92) (B:0.272)
30.0-39.9	2.99 (2.27-3.96) ($p=0.000$) (B:1.097)	2.09 (1.51-2.90) (B:0.739)†	1.97 (1.38-2.80) ($p=0.000$) (B:0.675)	1.44 (0.97-2.14) (B:0.364)
>40.0	12.62 (6.47-24.62) ($p=0.000$) (B:2.535)	7.27 (3.56-14.86) (B:1.984)†	2.93 (1.38-6.23) ($p=0.005$) (B:1.074)	1.62 (0.72-3.66) (B:0.481)
Smoking status				
Non-smoker	1	1	1	1
Ex-smoker	3.58 (2.62-4.90) ($p=0.000$) (B:1.275)	2.36 (1.64-3.38) (B:0.857)†	1.84 (1.28-2.63) ($p=0.001$) (B:0.608)	1.20 (0.81-1.79) (B:0.183)
Current smoking	3.18 (2.11-4.80) (B:1.157)	1.89 (1.19-3.01) (B:0.638)†	2.64 (1.66-4.20) ($p=0.000$) (B:0.972)	1.54 (0.92-2.56) (B:0.431)
Alcohol consumption				
No	1	1	1	1
Yes	2.88 (1.57-5.28) (B:1.058)	1.54 (0.78-3.06) (B:0.434)	3.46 (1.40-8.54) ($p=0.007$) (B:1.240)	3.2 (1.14-9.08) (B:1.169)
Major SDB symptoms				
No symptoms	1	1	1	1
Snoring (only)	1.73 (1.41-2.13) ($p=0.000$) (B:0.326)	1.41 (1.10-1.81) (B:0.347)*	1.39 (1.06-1.81) ($p=0.016$) (B:0.549)	1.06 (0.79-1.43) (B:0.059)
Witnessed apnea (only)	1.68 (1.24-2.26) ($p=0.001$) (B:0.383)	1.70 (1.20-2.41) (B:0.530)*	1.47 (1.00-2.14) ($p=0.047$) (B:0.516)	1.32 (0.87-1.99) (B:0.275)
Excessive day-time sleepiness (only)	1.07 (0.85-1.34) (B:0.343)	1.12 (0.87-1.47) (B:0.116)	1.41 (1.07-1.86) ($p=0.016$) (B:0.067)	1.50 (1.11-2.04) (B:0.408)†

The data are presented as adjusted odds ratios (95% confidence interval)

SDB: Sleep disordered breathing

* $p < 0.010$, † $p < 0.001$, ‡ $p < 0.050$

nessed apnea was most prevalent in 25-44 years age group and excessive daytime sleeping was most prevalent in >75 years age group (Figure 1). When the subjects were classified according to BMI, it was seen that all major symptoms related to SDB significantly increase as BMI increases (Figure 2).

DISCUSSION

The importance of this study is that it is the first population based study representing a city in Turkey. Although there are studies investigating SDB symptoms in truck, bus and taxi drivers in Turkey,^{26,27} to our knowledge, there are no studies investigating a sample population representing the whole city.

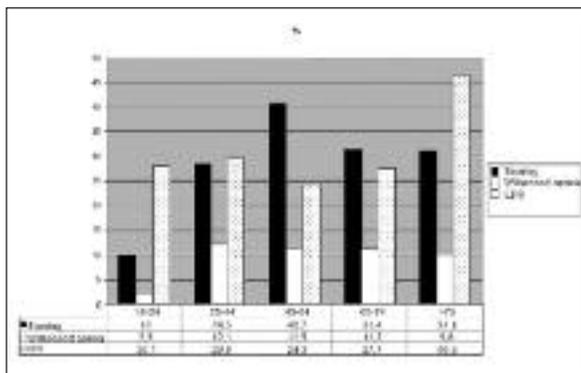


FIGURE 1: Major SDB symptoms according to age.

*In the comparison of the age groups p values for snoring, witnessed apnea and excessive daytime sleeping were $p < 0.001$, $p < 0.001$ and $p = 0.006$, respectively. EDS: Excessive daytime sleepiness.

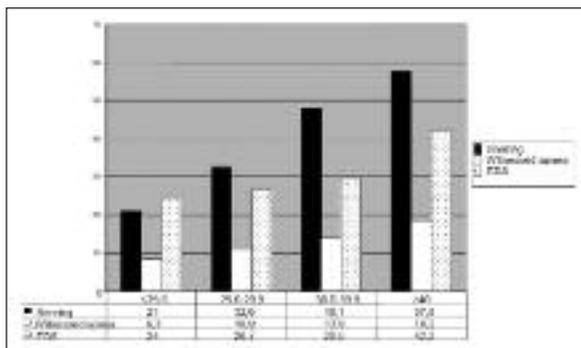


FIGURE 2: Major SDB symptoms according to BMI.

*In the comparison of the age BMI groups p values for snoring, witnessed apnea and excessive daytime sleeping were $p < 0.001$, $p = 0.010$ and $p = 0.020$, respectively. EDS: Excessive daytime sleepiness.

Prevalences of symptoms related with SDB varies in different studies. In a recent study from Greece, prevalence of habitual snoring was found 29.5% in males and 24.1% in females.²⁸ Mehra et al²⁹ found an overall prevalence of snoring as 29.9%. The rates in a Brazilian study by Pires et al³⁰ were 40% for males and 24% for females. Our results are similar to these studies. However, in a study from Spain, Marin et al³¹ reported a prevalence of loud snoring as 64% in males and 36% in females. They highlighted that using subjective reports, the prevalence of habitual snoring, may previously have been underestimated, as it was said to range from 30 to 45% in men. When comparing self and spouse reports of snoring, were compared poor agreement was found³² and hence in epidemiological studies on sleep disturbances, the bed-partner questionnaire or a direct measurement during sleep is mandatory. Using these approaches habitual snoring was detected in 64-81% of subjects in community studies.^{1,2,15}

In patients with OSAS, obesity is responsible for an increase in neck size; this so-called ‘neck obesity’ has been implicated in the pathophysiology of upper airway obstruction during sleep due to fat compression of the pharynx.^{33,34} In our study, all major symptoms related with SDB were significantly higher in the obese population. The high prevalences of witnessed apnea and excessive daytime sleepiness in women in our study was considered to be due to their obesity, since the women were more obese than men in our sample. On the other hand, some studies like Wisconsin Sleep Cohort Study³⁵ reported that white women with SDB had similar symptoms as those of men.

In this study, we also evaluated the relation of SDB symptoms with the concomitant diseases. DM, HT, hyperlipidemia and CHF were more prevalent among snorers. In epidemiologic studies, there is a growing consensus that SDB is an important risk factor for hypertension regardless of the excessive weight and other potentially confounding factors. Lindberg et al²⁰ reported that persistent snoring was an independent risk factor for the development of hypertension among males aged <50 years. Gislason et al²¹ showed that there was almost a three-

fold increase in the predicted prevalence of hypertension among intermittent and habitual snorer women when compared to nonsnorer women in the 40 to 49 years old age group and a 60 percent increase in the 50 to 59-year-old age group. Carlson et al³⁶ reported that age, BMI, and sleep apnea were all found to be independent predictors of hypertension. Obesity combined with sleep apnea turned out to result in a 3.9-fold increase in the prevalence of hypertension. In this study, we found a 1.7 folds increased HT risk in snorers.

Shin et al³⁷ reported that habitual snorers had significantly higher glucose and insulin levels at 2nd h of oral glucose tolerance test as compared to non-habitual snorers, while fasting glucose or insulin levels did not differ significantly between the groups. In a population-based sample of hypertensive men, the prevalence of obstructive sleep apnea was significantly higher in diabetic patients than in normoglycemic subjects, independent of central obesity.¹⁷ In contrast, in an Italian study of obese patients, snoring was associated with diabetes in a univariate analysis, however in a multivariate analysis, an independent effect of snoring was only observed for hypertension.³⁸

Elmasry et al²³ showed that in males aged 30-69 years, habitual snoring was associated with an increased incidence of diabetes within 10 years. They say that although obesity is the main risk factor for developing diabetes, coexistent habitual snoring may add to this hazard. Meslier et al¹⁸ reported a significant relationship between sleep-disordered breathing and impaired glucose-insulin metabolism that was independent of obesity and age. Al-Delaimy et al²⁴ also showed that snoring is independently associated with elevated risk of type II diabetes. Similar to these studies, we found a 1.4 folds increased DM risk in snorers.

The pathophysiology of hypertension and diabetes in sleep-disordered breathing is not fully understood. In a review of the pathophysiology of

hypertension in obstructive sleep apnea, Richert et al³⁹ stated that there were three components of importance: (1) large negative intrathoracic pressure, (2) intermittent hypoxemia and (3) arousal from sleep. As the severity of sleep apnea, as indicated by the maximum intraesophageal pressure, decreases with age,⁴⁰ this might be a possible explanation of why younger patients with sleep apnea are more prone to develop hypertension. When it comes to diabetes and impaired glucose metabolism, it is known that sleep deprivation reduces insulin sensitivity⁴¹ and has a harmful impact on carbohydrate metabolism similar to that seen in normal aging.^{42,43} The frequent arousals seen in patients with OSAS might have negative consequences on carbohydrate metabolism through similar pathophysiologic mechanisms, such as sleep deprivation or sleep disruption with other causes.

We also found a 1.4 folds increased risk for hyperlipidemia and 2.1 folds increased risk for CHF in our snorer population.

Snoring and witnessed apnea were found as risk factors for hypertension independently from smoking status, BMI or age. Excessive daytime sleepiness was an independent risk factor for diabetes mellitus.

A limitation for this study can be that it was a self-reported study which has the possibility of exaggeration or recall bias. To overcome this problem, we tried to investigate the medical reports and previous prescriptions of the subjects to verify the presence of the physician diagnosed diseases.

As a conclusion, the frequency of SDB symptoms were high in Afyon city, with a greater than expected rate for females. The risk for HT and DM were found to be higher in snorers. Snoring and witnessed apnea were found as risk factors for hypertension independently from smoking status, BMI or age, whereas excessive daytime sleepiness was an independent risk factor for diabetes mellitus.

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