



Correspondence to:

Sohrab Amiri
Medicine, Quran and Hadith Research Center
Baqiyatallah University of Medical Sciences,
Tehran, Iran
e-mail: Amirysohrab@yahoo.com

Sohrab Amiri 

Medicine, Quran and Hadith Research Center, Baqiyatallah University of Medical
Sciences, Tehran, Iran

Submitted: 02.12.2021
Accepted: 08.04.2023

Body mass index and sleep disturbances: a systematic review and meta-analysis

Abstract

Purpose: Overweight and obesity have been common non-communicable diseases in recent years that have far-reaching health effects. This study aimed to study the risk of sleep disturbances based on overweight and obesity using a systematic review and meta-analysis.

Views: Three sources of scientific information, namely PubMed, Scopus, and Embase, were selected as scientific databases and searched for articles published in English until May 2021. Also, a manual search on related articles was done. The resulting data required a meta-analysis based on odds ratios, which were extracted and then pooled. The odds ratio was calculated based on the random-effects method. Publication bias and the degree of heterogeneity were studied. Twenty-one longitudinal studies were selected for this meta-analysis. The odds ratio (OR) in this association between body mass index (BMI) ≥ 25 and sleep disturbances was 1.33 and the confidence interval (CI) was between 1.16-1.51. This OR was 1.12 and CI was between 0.98-1.29 in overweight. The OR was 1.24 and CI was between 1.07-1.44 in obesity. The OR was 1.92 and CI was between 0.89-4.15 in men. The OR was 1.47 and CI was between 1.15-1.89 in women. Publication bias and heterogeneity were evident in the meta-analysis.

Conclusions: Obesity increases the risk of sleep disturbances in the long term, and both obesity and sleep disturbances have negative effects on health, so it is necessary to develop appropriate interventions to improve obesity.

Key words: obesity, overweight, sleep disturbances, systematic review, meta-analysis.

INTRODUCTION

Obesity has become an epidemic in the world, with rates tripling between 1975 and 2016 [1]. Accordingly, in 1997 the World Health Organization recognized obesity as a pandemic that is a major cause of health problems [2]. The World Health Organization's 2016 report shows that 1.9 billion people over the age of 18 worldwide are overweight and the more than 650 million are obese; this means that 39% of people over the age of 18 are overweight and 13% are obese [3]. The Institute of Global Burden of Disease report also shows that the body mass index (BMI) of 25 kg/m² and above in adults between 1980 and 2013 in men increased from 28.8% to 36.9%, while the increase in women was from 29.8% to 38% [4]. There is also a significant increase in overweight and obesity among children and adolescents in both developed and developing countries [4]. In this regard, the increase in BMI in the form of overweight and obesity is associated with a range of diseases that are threats to health, such as mortality [5-7], cancer [8-11], diabetes [12-14], reduced quality of life [15-17], depression [18, 19], work-related

issues [20-22], and other mental health issues [23, 24]. One of the health dimensions that can be affected by overweight and obesity is sleep disturbances [25].

The International Classification of Sleep Disorders includes a variety of conditions [26]. The most prevalent type of sleep disorder is insomnia, with an estimated 30% of the adult population having trouble starting and continuing sleep, and between 5-10% having insomnia syndrome [27-29]. Another type of sleep disorder is obstructive sleep apnea, with a prevalence of 9% to 38% in the general population and is more prevalent in men [30]. It is estimated that among adults aged 30-69, 936 million worldwide have mild symptoms of obstructive sleep apnea and 425 million have moderate-to-severe obstructive sleep apnea [31]. Other types of sleep disorder can be noted as excessive daytime sleepiness; the overall prevalence of this disorder is reported to be 10.4% in men and 13.6% in women [32]. There are other types of sleep disturbance, such as poor sleep quality, sleep complaints, and so on. Several factors have been reported in connection with sleep disturbances, including aging [30, 33], sex, depressed mood [34], smoking [35], and alcohol use [36].

In addition, sleep disturbances have negative effects on various aspects of health and work such as increasing the risk of sick leave [37], dementia [38], cardiovascular disease [39], back pain [40], physical impairment [41], and mental disorders [42-44].

Studies have examined the risk factors associated with sleep disturbances [45-49]. The relationship between these and BMI and has been investigated in recent studies [30, 49, 50]. Accordingly, systematic review and meta-analysis studies have shown that BMI and weight loss are associated with a range of sleep disturbances, including the diagnosis and symptoms of insomnia [51], obstructive sleep apnoea [30, 52], and restless legs syndrome [53]. It is clear that overweight and obesity have widespread negative effects on various dimensions of health, and one of these dimensions is sleep. Three systematic reviews and meta-analyses have examined the relationship between BMI and sleep disturbances [30, 51-53]; however, reviewing this group of studies reveals several critical points and necessitates the need for a new systematic review and meta-analysis. First, out of the total number of these studies, only three [49, 51, 53] have meta-analyzed the relationship between BMI and sleep disturbances. Second, each of these three studies addressed only one type of sleep disturbance and did not provide a comprehensive meta-analysis of the association between BMI and sleep disturbances. Third, these studies of the results of research often relied on cross-sectional studies, although cross-sectional studies are not able to determine the causality between BMI and sleep disturbances. Although some studies have examined sex differences in the relationship between BMI and sleep disturbances, this has not been the case for a variety of other causes of sleep disturbances.

Based on what has been stated, this study sought to examine the risk of sleep disturbances in overweight and obesity based on longitudinal studies. This association was also examined separately for overweight and obesity. The third goal was to examine sex differences in the association between BMI and sleep disturbances. Finally, associations between BMI and three types of sleep disorders including insomnia, obstructive sleep apnea, and excessive daytime sleepiness were investigated.

METHODS

Search strategy

In this project, to guide the systematic review and meta-analysis process the steps mentioned in the PRISMA [54] protocol were used. Three sources of scientific information, namely PubMed, Scopus, and Embase, were selected as scientific databases, and based on a complete syntax of keywords listed in Appendix 1 each of these sources of information was fully searched; articles pub-

lished in English until May 2021 were included. A manual search on references in related articles was also done. In cases where the articles were ambiguous, the authors were contacted via email to provide the required information.

Selection criteria

A set of inclusion and exclusion criteria were used for this study. Studies were included in the project if they met the following requirements: 1) overweight and obesity were considered as exposure variables; 2) sleep disturbances such as insomnia, restless legs syndrome, obstructive sleep apnea, snoring, sleep complaints, poor sleep quality, and excessive daytime sleepiness were the outcome variables; 3) study design was longitudinal studies, and cross-sectional, retrospective, time series, clinical trial studies, and case studies – review studies were excluded. Studies were excluded from the project with one of the following conditions: 1) studies in which the study subjects were children; 2) studies that did not provide the necessary information for the odds ratio and making this calculation was not possible; 3) studies on women during pregnancy or following childbirth; 4) studies that were from the same database and in this case only one article with the highest quality was selected.

Data extraction

A framework for the information extraction process was created. Accordingly, types of information were extracted for each of the eligible studies and these included information sets including the authors of each study and year of publication, population, country, length of time until follow-up, demographic indicators of the study population such as age and sex, BMI, sleep disturbance and its measurement, and finally the result of each study.

Qualitative measure

In the qualitative evaluation section, as an essential component of a systematic review, the EPHPP tool [55, 56] was used. In the present study, according to the nature of this research, five adjusted dimensions of this tool were used to assess the quality of each of the studies selected.

Meta-analysis

This research involved a systematic review as well as a statistical analysis in the form of meta-analysis; the steps taken to complete the meta-analysis are described below. The odds ratio and confidence interval were extracted for each study. In cases where the risk ratio was reported, this ratio was converted into the odds ratio [57]. In cases where the hazard ratio was reported, this ratio was pooled separately. In some studies, there were dependent outcomes that used existing procedures [58] to combine them, and in independent outcomes, a fixed-effects meth-

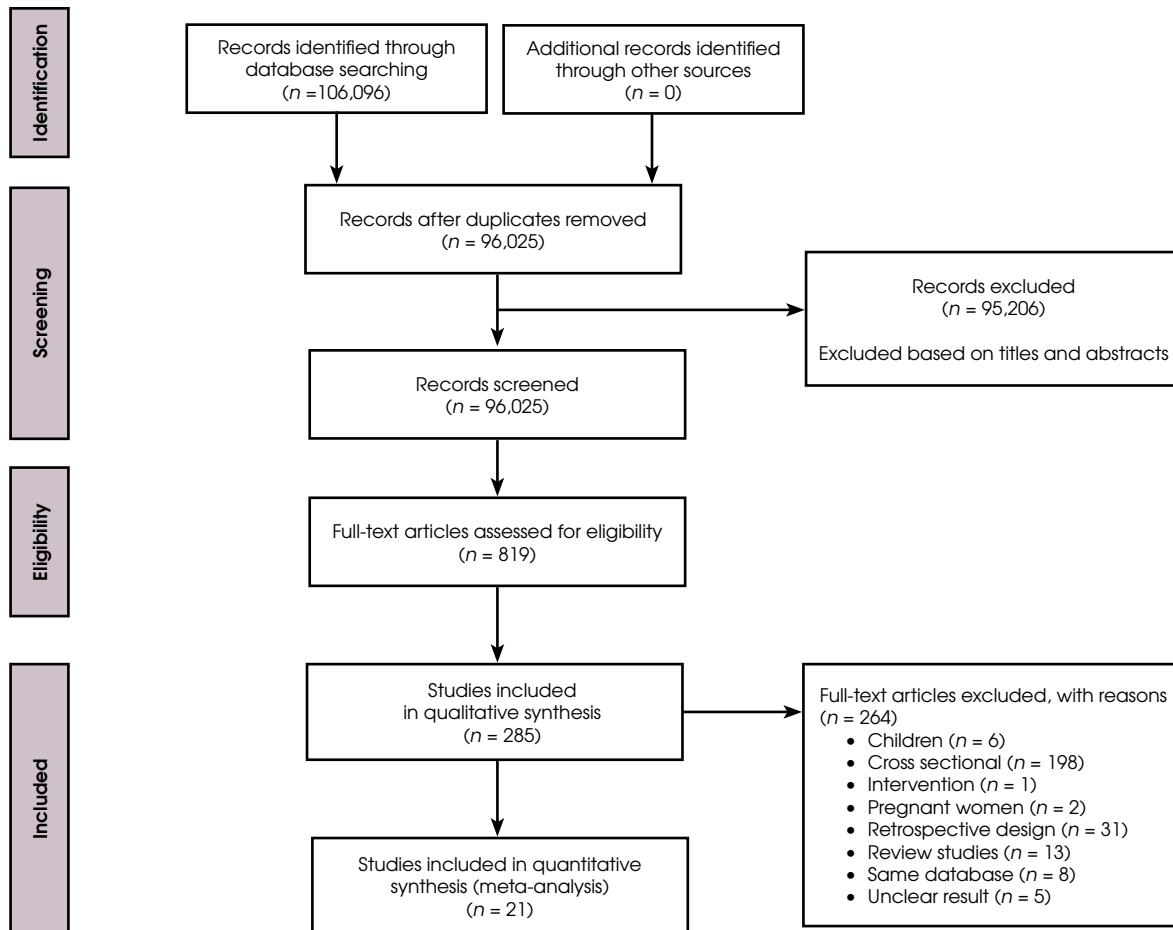


Figure 1. Selection flow diagram

od was used to combine them. Therefore, this information was extracted from each study and then the pooled odds ratio was calculated on the basis of the random-effects method. The analyzes were conducted at several levels. The first analysis was the association between BMI ≥ 25 and sleep disturbances, and then this odd ratio was calculated based on overweight and obesity, sex, and sleep disturbances categories. The degree of heterogeneity in the meta-analysis was studied based on two tests and their results were reported; the first test is I^2 , and the results of this test were reported for all analyzes, and the second was χ^2 test [59, 60]. Publication bias was investigated using tests (Egger [61], the trim-and-fill [62]), as well as graphical representations (Funnel plot). The results were analyzed with Stata-14 (Stata Corp. College Station, TX).

RESULTS

Selected studies

After searching the sources, 17,582, 27,315, and 61,199 articles were obtained from each of the information plat-

forms PubMed, Scopus, and Embase respectively. After screening the studies based on Figure I and removing the duplicate studies and the unqualified studies based on the titles and abstracts, 819 articles were downloaded and collected in full text. After reading all these full texts, 534 articles were excluded. Out of the remaining 285 articles, 264 were removed on the basis of the reasons mentioned in Figure I, and finally 21 longitudinal studies [63-83] remained for this project.

Quality of selected studies

Table 1 describes the qualitative status of studies in five dimensions: selection bias, confounder's bias, performance bias, data collection bias, and withdrawals/drop-outs and missing data bias. For selection bias, the results showed that a total of 14 studies had a low bias, while 3 articles were moderate in bias and the rest displayed high bias. When it came to confounder's bias, the frequency of high bias studies was higher. In terms of performance bias, all of the studies had a low and moderate bias. In the data collection method, six studies showed that the bias was low, and the rest showed moderate and

Table 1. Studies included in the meta-analysis

First author and year of publication	Country	Population	Follow-up time	Age range	Sex	Sample size	Body mass index	Sleep-related issues	Quality assessment: Risk of bias					Results	Adjustment for other covariates
									Selection	Confounders	Performance	Data collection method	Withdrawals and dropouts		
Batool-Anwar, 2016	USA	General population	6 years	25-75	76.04%	55,540	Self-reported	International Restless Legs Study Group criteria (restless legs syndrome)	Low	Low	Mode-rate	Low	Low	<p>Odd ratio; reference (BMI < 23)</p> <p>Men: 1. Overweight, BMI: 25-26.9; 1.41 (CI: 1.00-1.97); BMI: 27-29.9; 1.54 (CI 1.10-2.20); 2. Obesity, BMI > 30; 1.46 (CI: 0.96-2.20)</p> <p>Women: 1. Overweight, BMI: 25-26.9; 1.12 (CI: 0.91-1.38); BMI: 27-29.9; 1.66 (CI: 1.37-2.01); 2. Obesity, > 30; 1.64 (CI: 1.37-1.96)</p> <p>Pooled BMI ≥ 25: 1.48 (CI: 1.34-1.63)</p> <p>Pooled overweight: 1.41 (CI: 1.25-1.59)</p> <p>Pooled obesity: 1.61 (CI: 1.37-1.90)</p> <p>Pooled BMI ≥ 25 for men: 1.47 (CI: 1.19-1.81)</p> <p>Pooled BMI ≥ 25 for women: 1.48 (CI: 1.32-1.65)</p>	Race, the Crown-Crisp anxiety score, antidepressant medication, use of iron specific supplement at baseline, menopausal status, chronic illnesses
Bouloukaki, 2011	Greek	General population	6 years	50.7 ± 12.2	23.4% women	2,690	Measured	Apnea + hypopnea index (obstructive sleep apnea)	Low	High	Low	Low	Low	<p>Odd ratio</p> <p>Total*: 5.08 (CI: 4.37-5.91)</p> <p>Men: 6.86 (CI: 5.74-8.19)</p> <p>Women: 4.09 (CI: 3.01-5.54)</p>	Unadjusted
Chan, 2016	China	Coronary artery disease	3 years	> 21	9% women	587	Measured	Apnoea-hypopnoea index (obstructive sleep apnea)	Mode-rate	High	Low	Low	Low	<p>Odd ratio</p> <p>Reference: 1.11 (CI: 1.06-1.17)</p>	Smoking, hypertension
Chen, 2017	Taiwan	General population	7 years	≥ 18	46.6% women	12,728	Unknown	Clinically diagnosed (insomnia)	Low	Mode-rate	Mode-rate	Low	Low	<p>Hazard ratio</p> <p>Reference (BMI: 18.5-23.9)</p> <p>Overweight, BMI 24-26.9; 0.87 (CI: 0.73-1.03)</p> <p>Obesity, BMI ≥ 27; 0.99 (CI: 0.82-1.19)</p> <p>Pooled BMI ≥ 25: 0.92 (CI: 0.81-1.05)</p>	Multivariate adjustment
Elomaa, 2013	Finland	History of depressive symptoms	7 years	25-64	66.1% women	127	Measured	Self-reported (sleep disturbance)	High	High	Low	High	High	<p>Odd ratio</p> <p>Reference (without reference): 0.92 (CI: 0.84-1.00)</p>	Smoking, use of sleep or lipidlowering medication
Gureje, 2011	Nigeria	Elderly	1 year	≥ 65	50.5% women	1,307	Measured	World Health Organization Composite International Diagnostic Interview (insomnia)	Low	High	Low	Mode-rate	Mode-rate	<p>Odd ratio</p> <p>Reference: BMI < 18.5</p> <p>Overweight, BMI 25.0-29.9; 0.90 (CI: 0.30-2.70)</p> <p>Obesity, BMI ≥ 30: 1.70 (CI: 0.50-6.00)</p> <p>Pooled BMI ≥ 25: 1.19 (CI: 0.52-2.71)</p>	Sex, age
Janson, 2001	Sweden	General population	10 years	30-69	Men	2,602	Self-reported	Questionnaire (insomnia)	Mode-rate	Mode-rate	Mode-rate	Mode-rate	Mode-rate	<p>Odd ratio</p> <p>Reference (BMI < 27)</p> <p>BMI > 27: 1.35 (CI: 1.01-1.81)</p>	Medical disorder, BMI, physically inactive, alcohol dependent

Table 1. Cont.

First author and year of publication	Country	Population	Follow-up time	Age range	Sex	Sample size	Body mass index	Sleep-related issues	Quality assessment: Risk of bias					Results	Adjustment for other covariates
									Selection	Confounders	Performance bias	Data collection method	Withdrawals and dropouts		
Jaussef 2011	France	Older adults	Unknown	≥ 65	54.6% women	5,886	Self-reported	Questionnaire (sleep complaints) (insomnia)	Low	Moderate	Moderate	Moderate	Low	<p>Odd ratio</p> <p>Reference: BMI < 25</p> <p>Difficulty with initiating sleep</p> <p>Men: Overweight, BMI 25-29: 1.00 (CI: 0.51-1.96); Obesity, BMI > 30: 2.70 (CI: 1.15-6.34)</p> <p>Women: Overweight, BMI 25-29: 0.91 (CI: 0.61-1.37); Obesity, BMI > 30: 0.57 (CI: 0.30-1.07)</p> <p>Pooled BMI ≥ 25: 0.95 (CI: 0.71-1.27)</p> <p>Pooled overweight: 0.93 (CI: 0.66-1.32)</p> <p>Pooled obesity: 0.99 (CI: 0.60-1.65)</p> <p>Pooled BMI ≥ 25 for men: 1.46 (CI: 0.86-2.48)</p> <p>Pooled BMI ≥ 25 for women: 0.80 (CI: 0.57-1.12)</p>	Multivariate
Kim, 2004	Korea	Middle-aged	Unknown	40-69	32.3% women	457	Unknown	Apnea-hypopnea index (obstructive sleep apnea)	High	High	Moderate	Low	High	<p>Odd ratio</p> <p>Reference: 3.96 (CI: 2.27-6.91)</p>	Age, sex, hypertension
Knuiaman, 2006	Australia	General population	14 years	25-74	63% women	967	Measured	Questionnaire (snoring)	Low	Moderate	Low	Moderate	Low	<p>Odd ratio</p> <p>Reference: 1.44 (CI: 1.18-1.75)</p>	Multivariate-adjusted
Mokhlesi, 2012	USA	Polycystic ovary syndrome	Unknown	18-40	Women	78	Unknown	Berlin questionnaire (obstructive sleep apnea)	High	High	Moderate	Moderate	High	<p>Odd ratio</p> <p>1.23 (CI: 1.11-1.36)</p>	Unknown
Morgan, 2003	UK	Elderly	8 years	Unknown	Both	410	Unknown	Interview assessments (insomnia)	Moderate	Moderate	Moderate	High	High	<p>Odd ratio</p> <p>BMI > 25: 0.80 (CI: 0.50-1.40)</p>	Adjusted

Table 1. Cont.

First author and year of publication	Country	Population	Follow-up time	Age range	Sex	Sample size	Body mass index	Sleep-related issues	Quality assessment: Risk of bias					Adjustment for other covariates
									Selection	Confounders	Performance	Data collection	Withdrawals and dropouts	
Pedraza, 2012	USA	Elderly	3 years	≥ 75	62% women	1,085	Measured	Self-reported (sleep complaints, poor sleep quality)	Low	Low	Low	Mode-rate	Low	<p>Odd ratio; reference (BMI < 25)</p> <p>I. Trouble Falling</p> <p>1. Asleep: Overweight, BMI 25 < 30: 1.14 (CI: 0.72-1.81); Obesity, BMI 30 > 35: 1.11 (CI: 0.65-1.90); Over obesity, BMI ≥ 35: 1.73 (CI: 0.86-3.45)</p> <p>2. Wake up several times per night: Overweight, BMI 25 < 30: 1.11 (CI: 0.80-1.54); Obesity, BMI 30 > 35: 1.51 (CI: 1.03-2.21); Over obesity, BMI ≥ 35: 2.33 (CI: 1.37-3.96)</p> <p>II. Trouble Staying</p> <p>1. Asleep: Overweight, BMI 25 < 30: 1.02 (CI: 0.63-1.69); Obesity, BMI 30 > 35: 1.57 (CI: 0.92-2.68); Over obesity, BMI ≥ 35: 1.20 (CI: 0.55-2.62)</p> <p>2. Wake up after your usual amount of sleep. Feeling tired and worn: Overweight, BMI 25 < 30: 1.28 (CI: 0.73-2.22); Obesity, BMI 30 > 35: 1.58 (CI: 0.86-2.91); Over obesity, BMI ≥ 35: 1.91 (CI: 0.86-2.24)</p> <p>3. Poor quality of sleep: Overweight, BMI 25 < 30: 0.93 (CI: 0.61-1.41); Obesity, BMI 30 > 35: 1.25 (CI: 0.79-1.99); Over obesity, BMI ≥ 35: 1.20 (CI: 0.62-2.32)</p> <p>Pooled BMI ≥ 25: 1.32 (CI: 0.95-1.83) Pooled overweight: 1.53 (CI: 0.82-2.84) Pooled obesity: 1.16 (CI: 0.70-1.92)</p>
Phillips 2005	USA	General population	9 years	45-69	55.2% women	13,564	Unknown	Maastricht Questionnaire (sleep complaints)	Low	Low	Mode-rate	Mode-rate	Low	<p>Age, sex, menopausal status, race, education level, income, body mass index, depression, presence of cardiac disease, lung function status, presence of hypertension, alcohol intake, diabetes, and hypnotic use</p> <p>Odd ratio Reference: BMI 20-24 Difficulty falling asleep: Overweight, BMI 25-29: 0.80 (CI: 0.80-1.00); Obesity, BMI ≥ 30: 0.90 (CI: 0.80-1.01)</p> <p>Sleep continuity disturbance: Overweight, BMI 25-29: 1.00 (CI: 0.90-1.10); Obesity, BMI ≥ 30: 0.96-1.20 Non-restorative sleep: Overweight, BMI 25-29: 1.00 (CI: 0.90-1.10); Obesity, BMI ≥ 30: 1.00 (CI: 0.90-1.10) Pooled BMI ≥ 25: 0.97 (CI: 0.88-1.07) Pooled overweight: 0.93 (CI: 0.80-1.08) Pooled obesity: 1.00 (CI: 0.88-1.12)</p>
Quan, 2005	USA	Elderly	4 years	≥ 65	Both	4,467	Measured	Questionnaire (excessive daytime sleepiness)	Low	Mode-rate	Low	Mode-rate	Low	<p>Age, sex, race, time interval between examinations</p> <p>Odd ratio Reference: 1.03 (CI: 1.01-1.05)</p>

Table 1. Cont.

First author and year of publication	Country	Population	Follow-up time	Age range	Sex	Sample size	Body mass index	Sleep-related issues	Quality assessment: Risk of bias					Results	Adjustment for other covariates
									Selection	Confounders	Performance bias	Data collection method	Withdrawals and dropouts		
Singareddy, 2012	USA	General population	7.5 years	≥ 20	47.6% women	1,741	Self-reported	Self-reported (Insomnia)	Low	Low	Mode-rate	Mode-rate	Low	<p>Odd ratio</p> <p>Obesity: 1.24 (CI: 0.80-1.93)</p>	Age, gender, obesity, race, cigarettes, alcohol, caffeine use, physical health problems, mental health problems, poor sleep
Skarpsno, 2018	Norway	General population	11 years	≥ 20	54.5% women	21,847	Measured	Self-reported (Insomnia)	Low	Low	Mode-rate	Mode-rate	Low	<p>Risk ratio; reference (normal BMI)</p> <p>Men: Overweight: 1.22 (CI: 0.97-1.53), Obesity: 1.48 (CI: 1.08-2.01)</p> <p>Women: Overweight: 1.15 (CI: 0.97-1.35) Obesity: 1.25 (CI: 1.00-1.56)</p> <p>Odd ratio*</p> <p>Men: Overweight: 1.21 (CI: 0.97-1.50) Obesity: 1.45 (CI: 1.08-1.94)</p> <p>Women: Overweight: 1.14 (CI: 0.97-1.32) Obesity: 1.23 (CI: 1.00-1.51)</p> <p>Pooled BMI ≥ 25: 1.21 (CI: 1.09-1.34)</p> <p>Pooled overweight: 1.16 (CI: 1.03-1.32)</p> <p>Pooled obesity: 1.30 (CI: 1.10-1.54)</p> <p>Pooled BMI ≥ 25 for men: 1.29 (CI: 1.08-1.54)</p> <p>Pooled BMI ≥ 25 for women: 1.17 (CI: 1.04-1.33)</p>	Age, leisure time physical activity, education, smoking, shift work, alcohol consumption, HADS
Theorell-Haglöw, 2015	Sweden	General population	10 years	43.7 ± 15.2	Women	7,051	Self-reported	Uppsala Sleep Inventory; self-reported (excessive daytime sleepiness)	Low	High	Mode-rate	Mode-rate	Low	<p>Odd ratio</p> <p>Obesity, BMI ≥ 28: 1.28 (CI: 0.62-2.66)</p>	Age, development in all covariates, somatic disease
Tom, 2010	UK	General population	Unknown	48-54	Women	962	Unknown	Self-reported (sleep difficulty)	High	High	Mode-rate	High	Mode-rate	<p>Odd ratio, reference (BMI < 18.5)</p> <p>Overweight, BMI 25-29.9: 1.52 (CI: 0.80-2.89)</p> <p>Obesity, BMI ≥ 30: 1.82 (CI: 0.91-2.66)</p> <p>Pooled BMI ≥ 25: 1.69 (CI: 1.12-2.55)</p>	Age
Vahtera, 2009	France	Retired employees	7 years	35-50 at baseline	21% women	14,714	Self-reported	Questionnaires (sleep disturbances)	Low	High	Mode-rate	Mode-rate	Low	<p>Odd ratio, reference (BMI ≤ 24.9)</p> <p>Overweight, BMI 25-29.9: 1.03 (CI: 0.96-1.11)</p> <p>Obesity, BMI ≥ 30: 1.10 (CI: 0.97-1.24)</p> <p>Pooled BMI ≥ 25: 1.05 (CI: 0.98-1.12)</p>	Sex, age
Wilton, 2018	USA	Rheumatoid arthritis	≥ 14.8 years	≥ 18	68% women	813	Unknown	Medical record (obstructive sleep apnea)	Low	Mode-rate	Mode-rate	Low	Low	<p>Hazard ratio</p> <p>4.99 (CI: 2.98-8.36)</p>	Age, sex, calendar year

*Calculated by author(s)

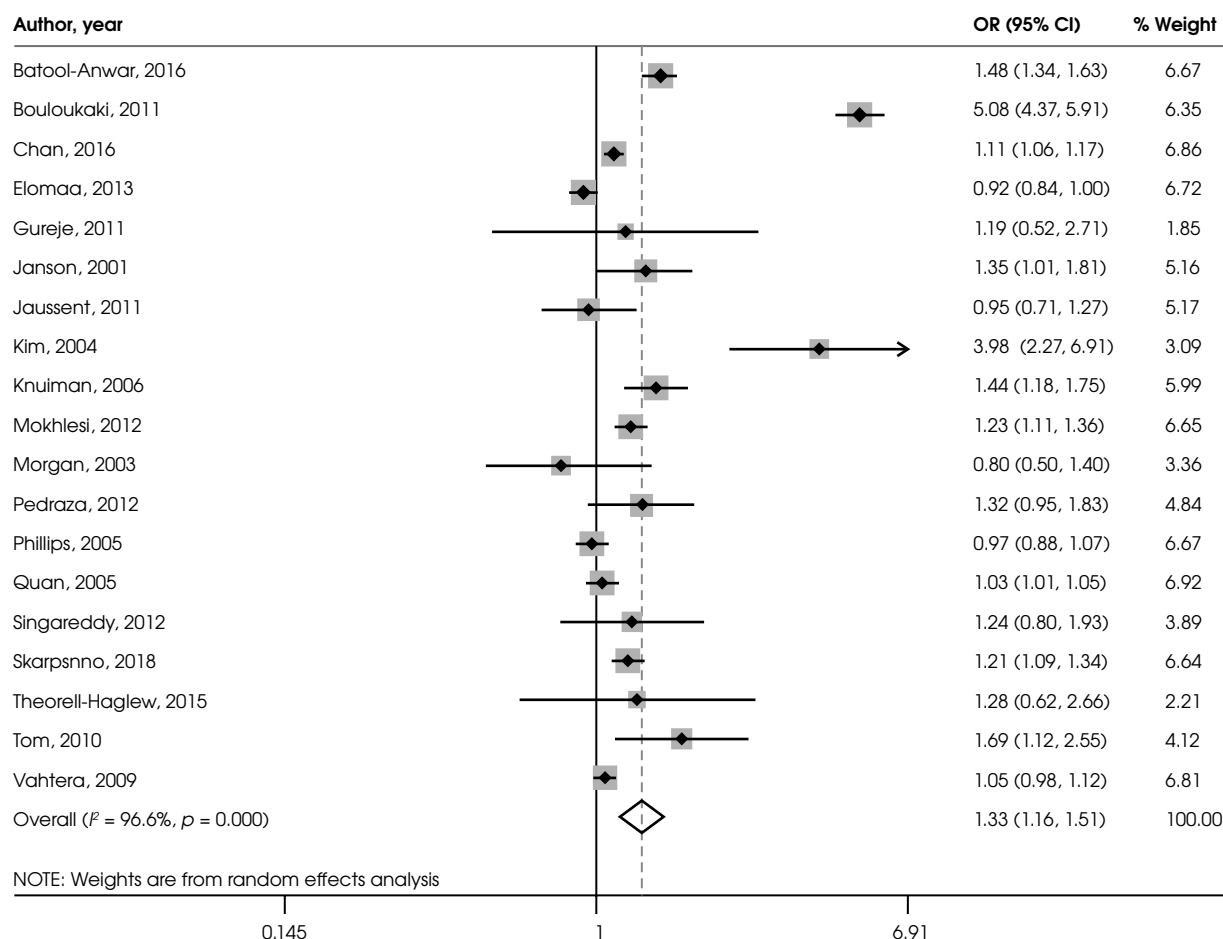


Figure II. The association between BMI ≥ 25 and sleep disturbances

high bias. In the component related to the withdrawals/dropouts and missing data, thirteen studies had a low bias, and the rest of the articles showed moderate and high bias.

Body mass index and risk of sleep disturbances

Figure II shows the association between BMI ≥ 25 and sleep disturbances. The odds ratio in this regard was 1.33 and the confidence interval was between 1.16-1.51 ($n = 19$; $p < 0.001$; $I^2 = 96.6\%$). For 2 studies that reported hazard ratio (HR), pooled HR was 2.10 and the confidence interval was between 0.40-11.01 ($n = 2$; $p = 0.379$; $I^2 = 97.4\%$).

Figure III shows the association between overweight and obesity and sleep disturbances. The odds ratio was 1.12 and the confidence interval was between 0.98-1.29 ($n = 8$; $p = 0.100$; $I^2 = 75.8\%$) in overweight. The odds ratio was 1.24 and the confidence interval was between 1.07-1.44 ($n = 10$; $p = 0.005$; $I^2 = 66.7\%$) in obesity.

Figure IV shows the association between BMI ≥ 25 with sleep disturbances in men and women. The odds ratio

was 1.92 and the confidence interval was between 0.89-4.15 ($n = 5$; $p = 0.095$; $I^2 = 98.2\%$) in men. The odds ratio was 1.47 and the confidence interval was between 1.15-1.89 ($n = 7$; $p = 0.002$; $I^2 = 91.7\%$) in women.

Figure V shows the association between BMI ≥ 25 and different types of sleep disturbance. The odds ratio was 2.23 and the confidence interval was between 1.17-4.26 ($n = 4$; $p = 0.015$; $I^2 = 99.2\%$) in obstructive sleep apnea. The odds ratio was 1.17 and the confidence interval was between 1.05-1.30 ($n = 6$; $p = 0.004$; $I^2 = 7.7\%$) in insomnia. The odds ratio was 1.03 and the confidence interval was between 1.01-1.05 ($n = 2$; $p = 0.003$; $I^2 = 0\%$) in excessive daytime sleepiness.

Heterogeneity and publication bias

The results of the heterogeneity test showed that this index is at a high level [59] and equal to $I^2 = 96.6\%$, and the χ^2 test showed this result 537.23 ($df = 18$; $p < 0.001$) [31]. The funnel test in Figure VI showed that there was publication bias. The Egger ($p = 0.071$) test was also significant, indicating a publication bias [61]. The trim-and-fill [62]

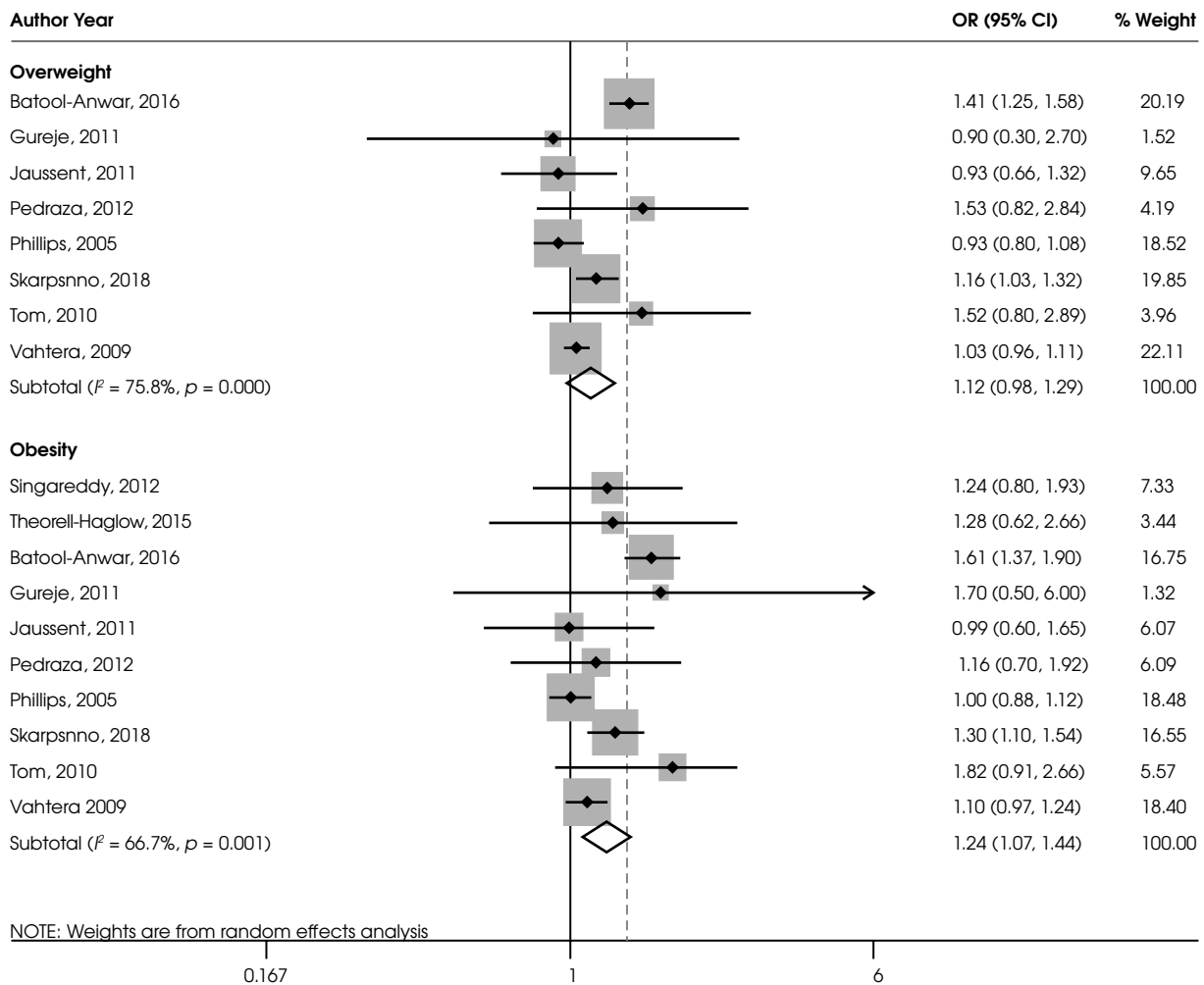


Figure III. The association between overweight and obesity and sleep disturbances

method imputed 6 missing studies, and the odds ratio increased to 1.59 and CI = 1.29-1.96 (Figure VI).

DISCUSSION

This study aimed to evaluate the risk of an individual experiencing sleep disturbances based on overweight and obesity, which was performed using systematic review and a meta-analysis of longitudinal meta-analyses. The first result of this meta-analysis was that a BMI above 25 increases the risk of sleep disturbances by up to 33%. This finding indicates the significant role that high BMI plays in the long term in increasing the risk of sleep disturbances. Also, discerning the relationship between overweight and obesity with the risk of sleep disturbances showed that with increasing BMI, the risk of sleep disturbances increases, so that obesity doubles the risk of sleep disturbances compared to overweight. There seems to be a direct linear relationship between BMI and sleep distur-

bances; as the BMI increases, so does the risk of sleep disturbances. This finding is consistent with previous studies that have shown that increasing BMI increases the risk of sleep disturbances [49, 53]. In the relationship between overweight and obesity and sleep disturbances, we need to pay attention to other mechanisms. One mechanism is the effect of overweight and obesity on the increased risk of diabetes [84, 85]; diabetes, in turn, is recognized as a risk factor for sleep disorders [86, 87].

Another finding of the study showed that the risk of sleep disturbances in men with overweight and obesity is almost twice as high compared to men with normal weight. In women with overweight and obesity, the risk of sleep disturbances was almost half that of men. This indicates that the risk of sleep disturbances is higher in men than in women based on overweight and obesity. A previous meta-analysis showed that women with obesity were more likely than men to suffer from restless legs syndrome [53]. It has also been shown that the prevalence

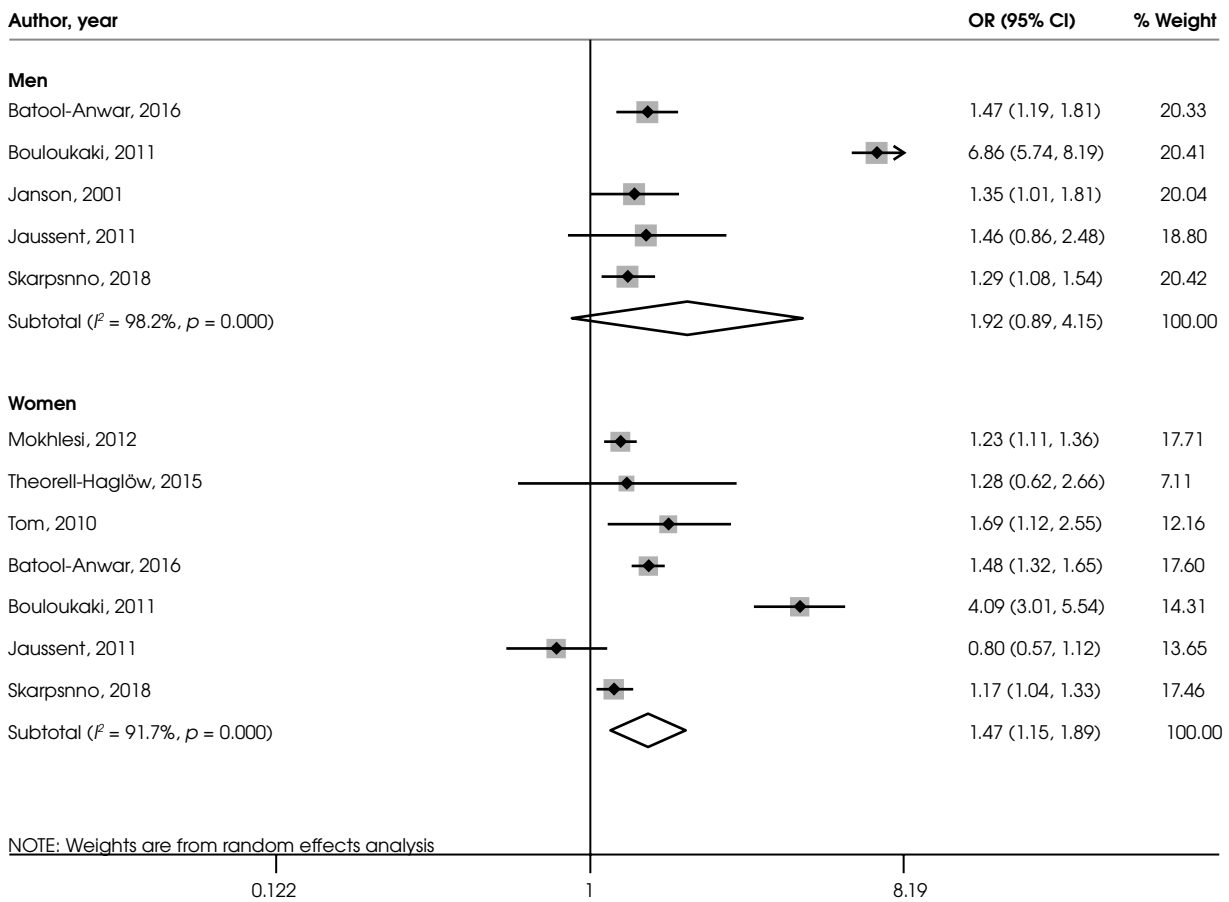


Figure IV. The association between BMI \geq 25 and sleep disturbances in men and women

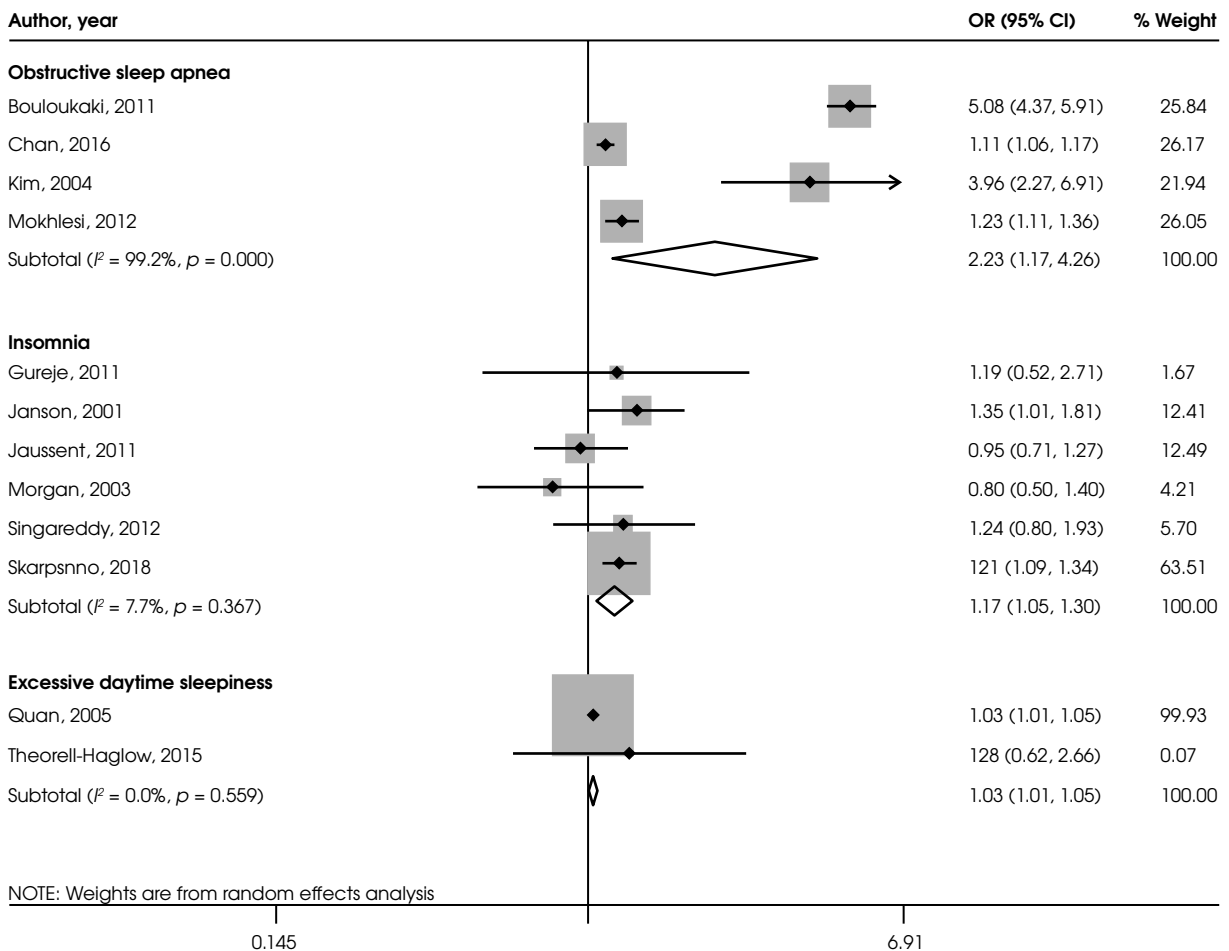


Figure V. The association between BMI \geq 25 and different types of sleep disturbance

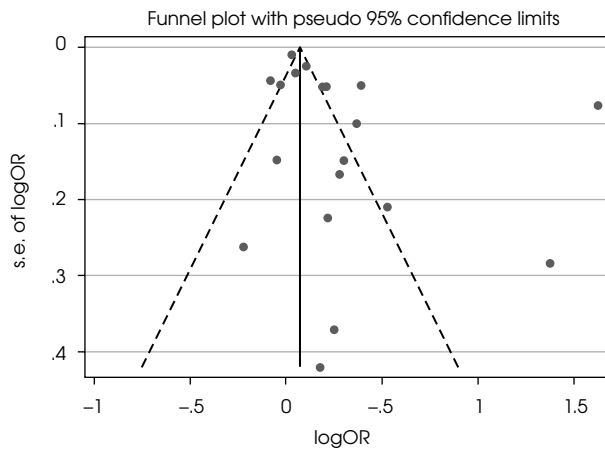


Figure VI. Funnel plot for BMI \geq 25 and sleep disturbances

of insomnia is higher in women than in men [88, 89]. What is clear is that in the current meta-analysis, the focus was also on obstructive sleep syndrome, so this difference in results could be due to the varying prevalence of different types of sleep disturbances in men and women, as studies have shown that obstructive sleep syndrome has a higher prevalence in men compared to women [90-93]. On the other hand, it should be noted that the current study examined sex differences in sleep disturbances based on BMI, and therefore the results may be different from those that have dealt with the prevalence of sleep disturbances in men and women.

Another finding on the relationship between BMI and types of sleep disturbance showed that overweight and obesity have the highest risk for obstructive sleep apnea

syndrome, doubling the risk of it. In the next ranks were insomnia and excessive daytime sleepiness. This high risk of obstructive sleep syndrome in people with obesity has been shown in studies to be up to three times higher than in normal-weight people [94-96]. Mechanisms through which obesity can increase the risk of obstructive sleep syndrome are listed [49], including an accumulation of fat in the respiratory tract and in the chest and abdomen that can cause problems with the respiratory system [49, 97, 98].

It should be noted that this study was able to provide a systematic review and meta-analysis of the association between BMI and sleep disturbances. On the other hand, overweight and obesity were studied separately, and also the results were studied based on sex and type of sleep disturbance. Due to the limitations of the existing literature, it was not possible to study other types of sleep disturbance. A methodological limitation was the high degree of heterogeneity in the studies included in the meta-analysis. Some studies included in the meta-analysis did not control confounder's variables well, which is a limitation.

CONCLUSIONS

The present study showed that with increasing BMI, the risk of sleep disturbances increases. Considering that the role of overweight and obesity in many health problems has been identified, paying attention to the impact of overweight and obesity on sleep disturbances can play a vital role in levels of prevention as well as treatment. Therefore, this finding could provide insights into how to reduce the existing levels of sleep disturbances.

Conflict of interest

Absent.

Financial support

Absent.

References

- Haththotuwa RN, Wijeyaratne CN, Senarath U. Worldwide epidemic of obesity. In: Mahmood TA, Arulkumaran S, Chervenak FA (eds.). *Obesity and Obstetrics*. 2nd ed. Elsevier; 2020, p. 3-8.
- World Health Organization. *Obesity: preventing and managing the global epidemic. Report of a WHO consultation*. Geneva: World Health Organization; 2000.
- World Health Organization. *Noncommunicable diseases country profiles 2018*. 2018; <https://www.who.int/news-room/fact-sheets/detail/obesity-and-overweight>.
- Ng M, Fleming T, Robinson M, et al. Global, regional, and national prevalence of overweight and obesity in children and adults during 1980-2013: a systematic analysis for the Global Burden of Disease Study 2013. *Lancet* 2014; 384: 766-781.
- Martínez J, Johnson CD, Sánchez-Payá J, de Madaria E, Robles-Díaz G, Pérez-Mateo M. Obesity is a definitive risk factor of severity and mortality in acute pancreatitis: an updated meta-analysis. *Pancreatology* 2006; 6: 206-209.

6. Flegal KM, Kit BK, Orpana H, Graubard BI. Association of all-cause mortality with overweight and obesity using standard body mass index categories: a systematic review and meta-analysis. *JAMA* 2013; 309: 71-82.
7. Aune D, Sen A, Prasad M, et al. BMI, and all-cause mortality: systematic review and non-linear dose-response meta-analysis of 230 cohort studies with 3.74 million deaths among 30.3 million participants. *BMJ* 2016; 353: i2156. DOI: 10.1136/bmj.i2156.
8. Moghaddam AA, Woodward M, Huxley RJCE, Biomarkers P. Obesity and risk of colorectal cancer: a meta-analysis of 31 studies with 70,000 events. *Cancer Epidemiol Biomarkers Prev* 2007; 16: 2533-2547.
9. Larsson SC, Wolk A. Obesity and the risk of gallbladder cancer: a meta-analysis. *Br J Cancer* 2007; 96: 1457-1461.
10. Larsson S, Wolk AJ. Overweight, obesity and risk of liver cancer: a meta-analysis of cohort studies. *Br J Cancer* 2007; 97: 1005-1008.
11. Yang Y, Dong J, Sun K, et al. Obesity and incidence of lung cancer: a meta-analysis. *Int J Cancer* 2013; 132: 1162-1169.
12. Guh DP, Zhang W, Bansback N, Amarsi Z, Birmingham CL, Anis AH. The incidence of co-morbidities related to obesity and overweight: a systematic review and meta-analysis. *BMC Public Health* 2009; 9: 88. DOI: 10.1186/1471-2458-9-88.
13. Abdullah A, Peeters A, de Courten M, Stoelwinder J. The magnitude of association between overweight and obesity and the risk of diabetes: a meta-analysis of prospective cohort studies. *Diabet Res Clin Pract* 2010; 89: 309-319.
14. Torloni MR, Betrán AP, Horta BL, et al. Prepregnancy BMI and the risk of gestational diabetes: a systematic review of the literature with meta-analysis. *Obesity Rev* 2009; 10: 194-203.
15. Pimenta FBC, Bertrand E, Mograbi DC, Shinohara H, Landeira-Fernandez J. The relationship between obesity and quality of life in Brazilian adults. *Front Psychol* 2015; 6: 966. DOI: 10.3389/fpsyg.2015.00966.
16. Ul-Haq Z, Mackay DF, Fenwick E, Pell JP. Meta-analysis of the association between body mass index and health-related quality of life among adults, assessed by the SF-36. *Obesity* 2013; 21: E322-E327. DOI: 10.1002/oby.20107.
17. Khairy SA, Eid SR, El Hadidy LM, Gebril OH, Megawer AS. The health-related quality of life in normal and obese children. *Egypt Pediatr Assoc Gazette* 2016; 64: 53-60.
18. Luppino FS, de Wit LM, Bouvy PF, et al. Overweight, obesity, and depression: a systematic review and meta-analysis of longitudinal studies. *Arch Gen Psychiatry* 2010; 67: 220-229.
19. Amiri S, Behnezhad S, Nadinlui KB. Body mass index (BMI) and risk of depression in adults: a systematic review and meta-analysis of longitudinal studies. *Obesity Med* 2018; 12: 1-12.
20. Sanchez Bustillos A, Vargas KG, Gomero-Cuadra R. Work productivity among adults with varied body mass index: results from a Canadian population-based survey. *J Epidemiol Global Health* 2015; 5: 191-199.
21. Ku B, Phillips KE, Fitzpatrick JJ. The relationship of body mass index (BMI) to job performance, absenteeism and risk of eating disorder among hospital-based nurses. *Appl Nurs Res* 2019; 49: 77-79.
22. Amiri S, Behnezhad SJ. Body mass index and risk of sick leave: a systematic review and meta-analysis. *Clin Obes* 2019; 9: e12334. DOI: 10.1111/cob.12334.
23. Amiri S, Behnezhad S. Obesity and anxiety symptoms: a systematic review and meta-analysis. *Neuropsychiatr* 2019; 33: 72-89.
24. Gariepy G, Nitka D, Schmitz N. The association between obesity and anxiety disorders in the population: a systematic review and meta-analysis. *Int J Obes (Lond)* 2010; 34: 407-419.
25. Chan WS, Levenson MP, McCrae C. A meta-analysis of associations between obesity and insomnia diagnosis and symptoms. *Sleep Med Rev* 2018; 40: 170-182.
26. American Academy of Sleep Medicine. *International Classification of Sleep Disorders*. 3rd ed. Darien, IL: American Academy of Sleep Medicine; 2014.
27. Morin CM, LeBlanc M, Daley M, Gregoire JP, Mérette C. Epidemiology of insomnia: prevalence, self-help treatments, consultations, and determinants of help-seeking behaviors. *Sleep Med* 2006; 7: 123-130.
28. Ohayon MM. Epidemiology of insomnia: what we know and what we still need to learn. *Sleep Med Rev* 2002; 6: 97-111.
29. Lichstein KL, Durrence HH, Taylor DJ, Bush AJ, Riedel BW. Quantitative criteria for insomnia. *Behav Res Ther* 2003; 41: 427-445.
30. Senaratna CV, Perret JL, Lodge CJ, et al. Prevalence of obstructive sleep apnea in the general population: a systematic review. *Sleep Med Rev* 2017; 34: 70-81.
31. Benjafield AV, Ayas NT, Eastwood PR, et al. Estimation of the global prevalence and burden of obstructive sleep apnoea: a literature-based analysis. *Lancet Respir Med* 2019; 7: 687-698.
32. Hayley AC, Williams LJ, Kennedy GA, Berk M, Brennan SL, Pasco JA. Prevalence of excessive daytime sleepiness in a sample of the Australian adult population. *Sleep Med* 2014; 15: 348-354.
33. Acquavella J, Mehra R, Bron M, Suomi JM, Hess GP. Prevalence of narcolepsy and other sleep disorders and frequency of diagnostic tests from 2013-2016 in insured patients actively seeking care. *J Clin Sleep Med* 2020; 16: 1255-1263.
34. Smagula SF, Stone KL, Fabio A, Cauley JA. Risk factors for sleep disturbances in older adults: evidence from prospective studies. *Sleep Med Rev* 2016; 25: 21-30.
35. Amiri S, Behnezhad S. Smoking and risk of sleep-related issues: a systematic review and meta-analysis of prospective studies. *Can J Public Health* 2020; 111: 775-786.
36. Britton A, Fat LN, Neligan A. The association between alcohol consumption and sleep disorders among older people in the general population. *Sci Rep* 2020; 10: 5275. DOI: 10.1038/s41598-020-62227-0.
37. Amiri S, Behnezhad S. Sleep disturbances and risk of sick leave: systematic review and meta-analysis. *Sleep Biol Rhythms* 2020; 18: 283-295.
38. Shi L, Chen SJ, Ma MY, et al. Sleep disturbances increase the risk of dementia: a systematic review and meta-analysis. *Sleep Med Rev* 2018; 40: 4-16.

39. Sofi F, Cesari F, Casini A, Macchi C, Abbate R, Gensini GF. Insomnia and risk of cardiovascular disease: a meta-analysis. *Eur J Prev Cardiol* 2020; 21: 57-64.
40. Amiri S, Behnezhad S. Sleep disturbances and back pain: systematic review and meta-analysis. *Neuropsychiatr* 2020; 34: 74-84.
41. Amiri S, Behnezhad S. Sleep disturbances and physical impairment: a systematic review and meta-analysis. *Phys Occup Ther Geriatr* 2021; 39: 258-281.
42. Harris LM, Huang X, Linthicum KP, Bryen CP, Ribeiro JD. Sleep disturbances as risk factors for suicidal thoughts and behaviours: a meta-analysis of longitudinal studies. *Sci Rep* 2020; 10: 13888. DOI: 10.1038/s41598-020-70866-6.
43. Pigeon WR, Pinquart M, Conner K. Meta-analysis of sleep disturbance and suicidal thoughts and behaviors. *J Clin Psychiatry* 2012; 73: e1160-e1167. DOI: 10.4088/JCP.11r07586.
44. Marino C, Andrade B, Campisi SC, et al. Association between disturbed sleep and depression in children and youths: a systematic review and meta-analysis of cohort studies. *JAMA Netw Open* 2021; 4: e212373. DOI: 10.1001/jamanetworkopen.2021.2373.
45. Smagula SF, Stone KL, Fabio A, Cauley JA. Risk factors for sleep disturbances in older adults: evidence from prospective studies. *Sleep Med Rev* 2016; 25: 21-30.
46. Zeng J, Wei M, Li T, et al. Risk of obstructive sleep apnea in Parkinson's disease: a meta-analysis. *PLoS One* 2013; 8: e82091. DOI: 10.1371/journal.pone.0082091.
47. Xiaolin Gu MM. Risk factors of sleep disorder after stroke: a meta-analysis. *Top Stroke Rehabil* 2017; 24: 34-40.
48. Bartel KA, Gradisar M, Williamson P. Protective and risk factors for adolescent sleep: a meta-analytic review. *Sleep Med Rev* 2015; 21: 72-85.
49. Dong Z, Xu X, Wang C, Cartledge S, Maddison R, Islam SMS. Association of overweight and obesity with obstructive sleep apnoea: a systematic review and meta-analysis. *Obesity Med* 2020; 17: 100185.
50. Andersen IG, Holm JC, Homøe PJ. Obstructive sleep apnea in obese children and adolescents, treatment methods and outcome of treatment – a systematic review. *Int J Pediatr Otorhinolaryngol* 2016; 87: 190-197.
51. Chan WS, Levens MP, McCrae CS. A meta-analysis of associations between obesity and insomnia diagnosis and symptoms. *Sleep Med Rev* 2018; 40: 170-182.
52. Wong AM, Barnes HN, Joosten SA, et al. The effect of surgical weight loss on obstructive sleep apnoea: a systematic review and meta-analysis. *Sleep Med Rev* 2018; 42: 85-99.
53. Lin S, Zhang H, Gao T, et al. The association between obesity and restless legs syndrome: a systemic review and meta-analysis of observational studies. *J Affect Disord* 2018; 235: 384-391.
54. Moher D, Liberati A, Tetzlaff J, Altman DG, The PG. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *PLoS Med* 2009; 6: e1000097. DOI: 10.1371/journal.pmed.1000097.
55. Armijo-Olivo S, Stiles CR, Hagen NA, Biondo PD, Cummings GG. Assessment of study quality for systematic reviews: a comparison of the Cochrane Collaboration Risk of Bias Tool and the Effective Public Health Practice Project Quality Assessment Tool: methodological research. *J Eval Clin Pract* 2012; 18: 12-18.
56. Thomas H. Quality assessment tool for quantitative studies. In: *Effective public health practice project*; 2003.
57. Zhang J, Yu KF. What's the relative risk? A method of correcting the odds ratio in cohort studies of common outcomes. *JAMA* 1998; 280: 1690-1691.
58. Borenstein M, Hedges LV, Higgins JP, Rothstein HR. *Multiple outcomes or time-points within a study. Introduction to meta-analysis.* John Wiley & Sons; 2009.
59. Higgins JP, Thompson SG. Quantifying heterogeneity in a meta-analysis. *Stat Med* 2002; 21: 1539-1558.
60. Ioannidis JP, Patsopoulos NA, Evangelou E. Uncertainty in heterogeneity estimates in meta-analyses. *BMJ* 2007; 335: 914-916.
61. Egger M, Smith GD, Schneider M, Minder CJB. Bias in meta-analysis detected by a simple, graphical test. *BMJ* 1997; 315: 629-634.
62. Duval S, Tweedie R. Trim and fill: a simple funnel-plot-based method of testing and adjusting for publication bias in meta-analysis. *Biometrics* 2000; 56: 455-463.
63. Batool-Anwar S, Li Y, De Vito K, Malhotra A, Winkelman J, Gao X. Lifestyle factors and risk of restless legs syndrome: prospective cohort study. *J Clin Sleep Med* 2016; 12: 187-194.
64. Bouloukaki I, Kapsimalis F, Mermigkis C, et al. Prediction of obstructive sleep apnea syndrome in a large Greek population. *Sleep Breath* 2011; 15: 657-664.
65. Chan PF, Tai BC, Loo G, et al. Optimal body mass index cut-offs for identification of patients with coronary artery disease at high risk of obstructive sleep apnoea. *Heart Lung Circ* 2016; 25: 847-854.
66. Chen LJ, Steptoe A, Chen YH, Ku PW, Lin CH. Physical activity, smoking, and the incidence of clinically diagnosed insomnia. *Sleep Med* 2017; 30: 189-194.
67. Elomaa AP, Koivumaa-Honkanen H, Niskanen L, et al. Self-reported sleep disturbance is associated with elevated levels of PAI-1 in individuals with a recorded history of depressive symptoms. *Prog Neuropsychopharmacol Biol Psychiatry* 2013; 47: 46-51.
68. Gureje O, Oladeji BD, Abiona T, Makanjuola V, Esan OJS. The natural history of insomnia in the Ibadan study of ageing. *Sleep* 2011; 34: 965-973.
69. Janson C, Lindberg E, Gislason T, Elmasry A, Boman GJS. Insomnia in men – a 10-year prospective population based study. *Sleep* 2001; 24: 425-430.
70. Jausseint I, Dauvilliers Y, Ancelin ML, et al. Insomnia symptoms in older adults: associated factors and gender differences. *Am J Geriatr Psychiatry* 2011; 19: 88-97.
71. Kim J, In K, Kim J, et al. Prevalence of sleep-disordered breathing in middle-aged Korean men and women. *Am J Respir Crit Care Med* 2004; 170: 1108-1113.

72. Knuiman M, James A, Divitini M, Bartholomew HJC. Longitudinal study of risk factors for habitual snoring in a general adult population: the Busselton Health Study. *Chest* 2006; 130: 1779-1783.
73. Mokhlesi B, Scoccia B, Mazzone T, Sam S. Risk of obstructive sleep apnea in obese and nonobese women with polycystic ovary syndrome and healthy reproductively normal women. *Fertil Steril* 2012; 97: 786-791.
74. Morgan K. Daytime activity and risk factors for late-life insomnia. *J Sleep Res* 2003; 12: 231-238.
75. Pedraza S, Al Snih S, Ottenbacher KJ, Markides KS, Raji MA. Sleep quality and sleep problems in Mexican Americans aged 75 and older. *Aging Clin Exp Res* 2012; 24: 391-397.
76. Phillips B, Mannino D. Correlates of sleep complaints in adults: the ARIC study. *J Clin Sleep Med* 2005; 1: 277-283.
77. Quan SF, Katz R, Olson J, et al. Factors associated with incidence and persistence of symptoms of disturbed sleep in an elderly cohort: the Cardiovascular Health Study. *Am J Med Sci* 2005; 329: 163-172.
78. Singareddy R, Vgontzas AN, Fernandez-Mendoza J, et al. Risk factors for incident chronic insomnia: a general population prospective study. *Sleep Med* 2012; 13: 346-353.
79. Skarpsno ES, Nilsen TI, Sand T, Hagen K, Mork P. Do physical activity and body mass index modify the association between chronic musculoskeletal pain and insomnia? Longitudinal data from the HUNT study, Norway. *J Sleep Res* 2018; 27: 32-39.
80. Theorell-Haglöw J, Åkerstedt T, Schwarz J, Lindberg EJS. Predictors for development of excessive daytime sleepiness in women: a population-based 10-year follow-up. *Sleep* 2015; 38: 1995-2003.
81. Tom SE, Kuh D, Guralnik JM, Mishra GJM. Self-reported sleep difficulty during the menopausal transition: results from a prospective cohort study. *Menopause* 2010; 17: 1128-1135.
82. Vahtera J, Westerlund H, Hall M, et al. Effect of retirement on sleep disturbances: the GAZEL prospective cohort study. *Sleep* 2009; 32: 1459-1466.
83. Wilton KM, Matteson EL, Crowson CS. Risk of obstructive sleep apnea and its association with cardiovascular and noncardiac vascular risk in patients with rheumatoid arthritis: a population-based study. *J Rheumatol* 2018; 45: 45-52.
84. Abdullah A, Peeters A, de Courten M, Stoelwinder J. The magnitude of association between overweight and obesity and the risk of diabetes: a meta-analysis of prospective cohort studies. *Diabet Res Clin Pract* 2010; 89: 309-319.
85. Bell JA, Kivimaki M, Hamer M. Metabolically healthy obesity and risk of incident type 2 diabetes: a meta-analysis of prospective cohort studies. *Obesity Rev* 2014; 15: 504-515.
86. Khalil M, Power N, Graham E, Deschênes SS, Schmitz N. The association between sleep and diabetes outcomes – a systematic review. *Diabet Res Clin Pract* 2020; 161: 108035. DOI: 10.1016/j.diabres.2020.108035.
87. Khandelwal D, Dutta D, Chittawar S, Kalra S. Sleep disorders in type 2 diabetes. *Indian J Endocrinol Metab* 2017; 21: 758-761.
88. Zeng LN, Zong QQ, Yang Y, et al. Gender difference in the prevalence of insomnia: a meta-analysis of observational studies. *Front Psychiatry* 2020; 11: 577429-577429.
89. Tsou MT. Gender-specific correlations of insomnia and attitudes toward treatment among community-dwelling elderly in Northern Taiwan. *Int J Gerontol* 2018; 12: 200-204.
90. Douglas NJ. Clinicians' guide to sleep medicine. Arnold; 2002.
91. Ancoli-Israel S, Kripke DF, Klauber MR, Mason WJ, Fell R, Kaplan O. Sleep-disordered breathing in community-dwelling elderly. *Sleep* 1991; 14: 486-495.
92. Young T, Palta M, Dempsey J, Skatrud J, Weber S, Badr S. The occurrence of sleep-disordered breathing among middle-aged adults. *N Engl J Med* 1993; 328: 1230-1235.
93. Quintana-Gallego E, Carmona-Bernal C, Capote F, et al. Gender differences in obstructive sleep apnea syndrome: a clinical study of 1166 patients. *Respir Med* 2004; 98: 984-989.
94. Barone JG, Hanson C, Dajusta DG, Gioia K, England SJ, Schneider D. Nocturnal enuresis and overweight are associated with obstructive sleep apnea. *Pediatrics* 2009; 124: e53-e59. DOI: 10.1542/peds.2008-2805.
95. Tishler PV, Larkin EK, Schluchter MD, Redline S. Incidence of sleep-disordered breathing in an urban adult population: the relative importance of risk factors in the development of sleep-disordered breathing. *JAMA* 2003; 289: 2230-2237.
96. Hanis CL, Redline S, Cade BE, et al. Beyond type 2 diabetes, obesity and hypertension: an axis including sleep apnea, left ventricular hypertrophy, endothelial dysfunction, and aortic stiffness among Mexican Americans in Starr County, Texas. *Cardiovasc Diabetol* 2016; 15: 86. DOI: 10.1186/s12933-016-0405-6.
97. Yaggi HK, Concato J, Kernan WN, Lichtman JH, Brass LM, Mohsenin V. Obstructive sleep apnea as a risk factor for stroke and death. *N Engl J Med* 2005; 353: 2034-2041.
98. Ramar K, Dort LC, Katz SG, et al. Clinical practice guideline for the treatment of obstructive sleep apnea and snoring with oral appliance therapy: an update for 2015. *J Clin Sleep Med* 2015; 11: 773-827.