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## Original Article

# Sleep quality does not mediate the negative effects of chronodisruption on body composition and metabolic syndrome in healthcare workers in Ecuador

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## ABSTRACT

**Background and aims:** The objective of the present work was to determine to what extent sleep quality may mediate the association between chronodisruption (CD) and metabolic syndrome (MS), and between CD and body composition (BC).

**Methodology:** Cross-sectional study which included 300 adult health workers, 150 of whom were night shift workers and thereby exposed to CD. Diagnosis of MS was made based on Adult Treatment Panel III criteria. Sleep quality was measured using the Pittsburgh Sleep Quality Index. Body mass index (BMI), fat mass percentage, and visceral fat percentage were measured as indicators of body composition (BC). Data were analyzed using logistic, linear regression and structural equation models.

**Results:** The odds of health workers exposed to CD to suffer MS was 22.13 (IC<sub>95</sub> 8.68–66.07) when the model was adjusted for age, gender, physical activity and energy consumption. CD was also significantly associated with an increase in fat mass and visceral fat percentages, but not to BMI. Surprisingly, there was not enough evidence supporting the hypothesis that sleep quality contributes to the association between CD and MS or between CD and BC.

**Conclusions:** Sleep quality does not mediate the negative effects of CD on MS nor on BC.

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## 1. Introduction

Chronodisruption (CD), which concerns a significant disturbance of the circadian organization of physiology, metabolism and behavior, can lead to disturbances in metabolic activity, potentially generating a negative vicious circle [1,2]. Indeed, lack of synchronization of internal biological systems relative to external, i.e., environmental changes, may have negative effects on health and

eventually result in chronic disturbances, including metabolic syndrome (MS) [2–7]. MS is characterized by co-occurrence of interrelated metabolic factors such as insulin resistance (diabetes type 2), abdominal obesity and atherogenic dyslipidemia [8,9]. As sleep has been recognized as one of the modulators of metabolic homeostasis, sleep disorders may form one of the risk factors contributing to MS [10,11].

CD can be artificially induced by working in night shifts, which involves non physiological rest schedules and exposure to artificial light at night [3]. In line with the association between CD and metabolic symptoms [2–7], night shift workers have an increased risk of suffering from not only MS, but also from cardiovascular diseases including coronary events, myocardial infarction and ischemic stroke [12–14]. Likewise, night shift workers, who also

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show increased odds of smoking and consuming higher amounts of caffeine [2,6,7], suffer from body weight increase [15], often are faced with severe sleep disturbances [10,11]. These findings raise the possibility that hampered quality of sleep is one of the main factors contributing to the link between CD and MS. To investigate this hypothesis we set out to investigate whether sleep quality mediates the association between CD and MS and between CD and BC in a group of night shift workers and a group of regular day workers for control.

## 2. Subjects, materials and methods

### 2.1. Subjects

A non-experimental cross-sectional design was implemented. A total of 300 volunteers who worked in health care centers in the central area of Ecuador were recruited. The sample comprised 150 night shift workers and 150 day workers, including adult men and women between 18 and 60 years old. A subject was considered a night shift worker if he or she was working 8 h or more during the night, from 7:00 p.m. to 06:00 a.m. (when the sun remains hidden in Ecuador), over the last six months. All data were collected by trained and qualified personnel.

### 2.2. Ethical considerations

The study was performed in accordance with the ethical guides of the Helsinki Declaration of Human Studies. The written informed consent was obtained from all the participants of the study. Patient data were codified to guarantee anonymity. The study was approved by a institutional review committee of the Research Institute of the Escuela Superior Politécnica de Chimborazo.

### 2.3. Measurements

#### 2.3.1. Anthropometry and body composition (BC)

Height was measured with a rigid and in-extensible wall tape of 60–210 cm with an accuracy resolution of 0.1 cm; weight was measured with a 50 gr precision scale; and waist circumference was measured with a 0.6 mm wide metal-tape taking the midpoint of the distance between the edge of the iliac crest and the ribcage. Subjects were measured with the least amount of clothing possible, 2–6 h post prandial. Body mass index (BMI) was calculated using the equation:  $BMI = \text{weight (kg)}/\text{height (m)}^2$ . Fat mass and visceral fat were measured with a electrical bioimpedance scale, brand SECA, with a reading range of 0–120 kg, and a precision of 100 g.

#### 2.3.2. Sleep quality

The Spanish version of the Pittsburgh Sleep Quality Index (PSQI) was used to determine sleep quality [16]. This index consists of 19 self-evaluated items, which are grouped into seven components, including subjective sleep quality, sleep latency, sleep duration, habitual sleep efficiency, presence of a sleep disorder, use of hypnotic medications, and daytime dysfunction. Each of these components receives a score of 0–3 points. The total score is the result of the sum of the seven components, so the test can give a total score of 21 points. Higher scores indicate worse quality of sleep.

#### 2.3.3. Caloric intake

Caloric intake was calculated based on the amount of food intake. To measure food intake, participants filled out a food diary for every day of the last week. Data were analyzed using the Ecuadorian food composition table, obtaining average amounts of caloric intake measured in kilo calories (kcal) for each day for each subject. Caloric intake was then calculated averaging kcal

consumed during the week. Prior registration, participants were trained by a nutritionist on how to fill out the diary. Participants were also provided with a telephone number for communication in case of doubts.

#### 2.3.4. Biochemical parameters

Triglycerides, HDL cholesterol and fasting glycaemia values were obtained from clinical records. These values were used only if measurements were made no more than 12 weeks before the other measurements were made.

#### 2.3.5. Systolic and diastolic blood pressure

A SECA sphygmomanometer was used to measure blood pressure (BP). Systolic blood pressure (SBP) and diastolic blood pressure (DBP) were measured in mm Hg following international recommendations for conventional, ambulatory and home blood pressure measurements [17]. BP was measured twice with a 5-min time lapse between each measurement. If the difference between measurements was  $\geq 5$  mm Hg, a third measurement was made 10 min after the subject kept rest.

#### 2.3.6. Metabolic syndrome (MS)

The Third Report of the Expert Panel on Detection, Evaluation, and Treatment of High Blood Cholesterol in Adults (also referred to as the Adult Treatment Panel III or ATP III criteria) was used to diagnose MS [8]. MS was diagnosed when three or more of the following criteria were met: 1) abdominal obesity (waist circumference in men  $> 102$  cm and in women  $> 88$  cm); 2) triglycerides  $\geq 50$  mg/dl; 3) HDL cholesterol (in men  $< 40$  mg/dl and in women  $< 50$  mg/dl); 4) BP  $\geq 130/85$  mm Hg; and 5) fasting glucose  $\geq 110$  mg/dl.

#### 2.3.7. Physical activity

The short version of the International Physical Activity Questionnaire (IPAQ) for adults was used to measure the physical activity of the subjects in the study sample [18]. The IPAQ provides information on the time spent walking, activities of moderate and vigorous intensity, and sedentary behavior during the 7 days prior to the survey. Accordingly, the subjects were classified into an active, moderately active, or sedentary group.

### 2.4. Statistical analyses

Results were expressed as means and standard deviations for quantitative variables with normal distribution and frequencies and percentages for categorical variables. Differences regarding age, gender, physical activity, sleep quality, energy consumption, body composition, ATP III criteria and MS were determined using  $\chi^2$  test, *t*-test, and Wilcoxon signed rank test, depending on the type of the variable (see Appendix A). To determine the association between MS and night shift work we implemented binomial logistic regression models, considering MS as a response variable and night shift work as an explanatory variable. Models were adjusted for age, gender, physical activity, and energy consumption. To determine the association between BMI, fat mass, visceral fat and shift work we implemented linear regression models, considering BMI, fat mass, and visceral fat as response variables and shift work as an explanatory variable. Models were adjusted for age, gender, physical activity, and energy consumption.

In order to find out whether sleep quality contributes to the association between night shift work, MS and BC we followed the causal steps approach outlined by Baron & Kenny in 1986 [19]. Mediation analysis allows the estimation of a direct effect relating the independent to the depended variable and a mediated effect (indirect) by which the independent variable indirectly affects the

dependent variable through the hypothesized mediating variable. We defined MS as the dependent variable, shift work as the independent variable and sleep quality as the hypothesized mediating variable. BC was modeled as the latent factor determined by BMI, fat mass and visceral fat as indicators. To perform mediation analysis, we used structural equation modeling (SEM). Mediation analysis was performed using Lavaan v0.6-5 [20]. All statistical analyses were performed using R v3.6.3 [21].

### 3. Results

#### 3.1. General characteristics of the sample

The sample included 300 adult men and women (65% and 35%, respectively), half of whom were exposed to CD, because they were night shift workers ( $n = 150$  vs.  $n = 150$ ). Mean age of people of the sample was 38.11 years (SD 7.83). Distribution of gender was similar among both groups of workers ( $p = 0.09$ ; Table 1; Appendix A). People who worked in shifts were older, more sedentary, and had a higher energy intake (kcal/day) than people who did not work in shifts (all  $p < 0.05$ ; Table 1; Appendix A). Shift workers also had a higher BMI, percentage of fat mass and visceral fat than those non-shift workers (all  $p < 0.05$ ; Table 1; Appendix A).

Moreover, distribution of MS was statistically significantly different among both groups of workers ( $p < 0.001$ ; Table 1; Appendix A). Percentage of people who met the ATP III criteria for MS was higher in the group of shift workers in comparison to non-shift workers (56% vs. 4%; Table 1). Table 1 shows the breakdown of MS criteria among workers. Except for DBP, mean values of waist circumference, glucose, SBP, and triglycerides of shift workers were statistically significantly higher than those of non-shift workers (all  $p < 0.05$ ; Table 1; Appendix A). Moreover, cholesterol HDL was significantly lower ( $p < 0.001$ ; Table 1).

#### 3.2. Both metabolic syndrome and body composition are associated with shift work

Health workers that were exposed to CD showed higher odds of MS in comparison to people who were not exposed to CD (OR = 30.55; CI<sub>95</sub> 13.70–81.64) (Table 2, non-adjusted model). This OR decreased to 22.13 (CI<sub>95</sub> 8.68–66.07) when the model was adjusted for age, gender, physical activity and calorie intake (Table 2, adjusted model). In order to study the relationship between BC and CD we assessed the association between BMI, percentage of fat mass and visceral fat as indicators of BC and night shift work. From the three indicators, only BMI did not show a statistically significant association with shift work when the model was adjusted for age, gender, physical activity and calorie intake (Table 2, adjusted model). Working in night shifts increased percentage of fat mass by 3.32% (95% CI 1.71:4.92) and percentage of visceral fat by 2.24% (95% CI 1.54:2.96).

#### 3.3. Sleep quality does not mediate the association between shift work and metabolic syndrome

Next, we investigated whether sleep quality could mediate the effect of CD on MS. Accordingly, we tested this hypothesis using SEM (see Methods). We observed that most of the effect of shift work on MS was transmitted directly and not by mean of sleep quality. For instance, from the total standardized total effect (0.58,  $p < 0.001$ ), all of it was transmitted directly (Table 3). Moreover, mediated effect (indirect effect) did not reach statistical significance (Table 3).

#### 3.4. Sleep quality does not mediate the association between shift work and body composition

Finally, we investigated whether sleep quality could mediate the effect of shift work on BC. Accordingly, we tested this hypothesis using SEM (see Methods). We modeled BC as a latent factor using as indicator BMI, fat mass and visceral fat as indicators. We observed that most of the effect of shift work on BC was transmitted directly and not by mean of sleep quality. For instance, from the total standardized total effect (0.36,  $p < 0.001$ ), 0.33 of it was transmitted directly (Table 3). Moreover, mediated effect (indirect effect) did not reach statistical significance (Table 3).

### 4. Discussion

The current study on health workers subjected to day-night shifts investigated the association between CD, MS, and BC, and evaluated whether sleep quality mediates their relationships. Our findings suggest that CD increases the odds of MS and changes BC in that, it increases the percentage of both fat mass and visceral fat. Interestingly, there was not sufficient evidence indicating that sleep quality mediates the deleterious impact of CD on health.

#### 4.1. Shift work and metabolic syndrome

We observed a significant association between CD and MS. The odds of MS of workers exposed to CD was near 22 times higher than the odds of workers who were not exposed to CD. Our findings are in agreement with those of two recent studies, a meta-analysis and a systematic review [22,23]. Moreover, the results of these studies suggest that the effect of CD on MS largely depends on the time in which a person has started to be exposed to night shift work; being more deleterious if it started earlier in life. Taken together, we can conclude that night shift work should be monitored, especially from the perspective of occupational risk and care of professionals exposed to CD. Furthermore, according to the latest available data from a national survey in Ecuador, more than 2 million people between 20 and 59 years of age (near 23%) suffer from MS [24]. Because MS constitutes a serious public health concern, identification of vulnerable groups such as night shift workers is a necessary contribution to Ecuadorian public health that requires special attention.

#### 4.2. Chronodisruption and body composition

Our study shows that people who are exposed to CD have habits generally associated with unhealthy body composition. In fact, they were more sedentary and showed a higher energy intake compared to day workers. These findings are in agreement with those of two recent studies suggesting that night shift workers require interventions to improve their life-style habits [25,26]. For example, nurses working in night shifts were found to have a higher energy consumption than required by their nutritional needs, causing excessive weight gain [25]. Yet, some other studies indicate that the relations described above might be more subtle or at least be subject to compensatory behavior.

Indeed, the study by Hulsege and colleagues found no significant differences in the frequency of food intake or quality of food among night shift and day workers, but they did find differences in the quality of snacks a person consumed depending on whether they worked during the day or at night [27]. People tended to consume healthier snacks when the work-shift occurred during daytime, compensating for the more unhealthy night shifts. These findings indicate that it is necessary to deeply understand eating habits of shift workers in order to understand the factors that control their body composition.

**Table 1**  
**General characteristics of the sample.** Sample included 300 people, 150 who worked in shifts and 150 who did not. General characteristics of the population are shown in relation to the presence or absence of shift work.

		Shift work							
		No (n = 150)				Yes (n = 150)			
		n	%	Mean	SD	n	%	Mean	SD
Age***				36.37	8.01			39.85	7.25
Gender <sup>NS</sup>	Male	60	40			46	31		
	Female	90	60			104	69		
Physical activity***	Active	38	25			1	1		
	MA	87	58			36	24		
	Sedentary	25	17			113	75		
PSQI***				7.23	1.55			11.93	3.96
Energy intake (kcal/d)***				2139.13	382.36			2327.56	526.91
Body composition	BMI (kg/m <sup>2</sup> )**			23.46	3.60			24.58	3.53
	Fat mass (%)***			29.49	5.53			35.08	5.97
	Visceral fat (%)***			12.19	2.41			15.18	2.54
ATP criteria	Waist circumference (cm)***			82.50	13.00			88.09	9.93
	Glucose (mg/dl)***			101.89	7.49			112.72	11.38
	DBP (mm Hg) <sup>NS</sup>			76.77	8.83			77.40	12.68
	SBP (mm Hg)***			112.03	8.35			118.53	11.87
	Cholesterol HDL (mg/dl)***			54.05	9.99			40.97	3.10
	Triglycerides (mg/dl)***			138.30	9.89			157.80	18.82
Metabolic syndrome***	No	144	96			66	44		
	Yes	6	4			84	56		

Abbreviations and nomenclature: n = number; kg = kilograms; kg/m<sup>2</sup> = kilograms per squared meter; kcal/d = kilocalories per day; SD = standard deviation; BMI = body mass index; % = percentage; MA = moderately active; PSQI = Pittsburgh Sleep Quality Index.

\* = p < 0.05; \*\* = p < 0.010; \*\*\* = p < 0.001; NS = not significant.

**Table 2**  
**Association between working shift and MS and body composition.** Model was adjusted for age, gender, physical activity and energy consumption.

Variables	Unadjusted model					Adjusted model				
	OR	β	95% IC		p	OR	β	95% IC		p
			LI	UI				LI	UI	
<b>MS (yes)</b>	30.55		13.70	81.64	<0.001	22.13		8.68	66.07	<0.001
<b>BMI (kg/m<sup>2</sup>)</b>		1.11	0.30	1.92	0.007		0.06	-0.91	1.03	0.899
<b>Fat mass (%)</b>		5.58	4.27	6.89	<0.001		3.32	1.71	4.92	<0.001
<b>Visceral fat (%)</b>		2.99	2.42	3.55	<0.001		2.24	1.54	2.96	<0.001

Abbreviations: OR = odds ratio; IC = confidence interval; LI = lower limit; UI = upper limit; p = p-value.

**Table 3**  
**Mediation analysis for MS and BC.** Tested mediator was PSQI. Model was adjusted for age, gender, physical activity and energy consumption. Model was fitted using SEM. Body composition was modeled as a latent factor using BMI, fat mass, and visceral fat.

Pathway	Standardized est.	p
<b>Mediation for MS</b>		
MS < - shift work (total effect)	0.58	<0.001
MS < - shift work (direct effect)	0.59	<0.001
MS < - PSQI (indirect effect)	-0.02	0.710
<b>Mediation for BC</b>		
BC < - shift work (total effect)	0.36	<0.001
BC < - shift work (direct effect)	0.33	<0.001
BC < - PSQI (indirect effect)	-0.03	0.325

Abbreviations: MS = metabolic syndrome; BC = body composition; PSQI = Pittsburgh Sleep Quality Index; p = p-value; <- = depends on.

Night shift workers in the current study also presented less healthy values with respect to various indicators of BC, which included BMI, percentage of fat mass and visceral fat. Classically, studies on the regulation of body weight have almost exclusively focused on caloric intake and energy expenditure. However, evidence is emerging that energy regulation may be linked to circadian clock disruption and CD, suggesting that the timing of food intake itself could play an important role in weight control [28]. This notion is supported by the observation that suppression of

melatonin, which occurs during shift work, induces insulin resistance, glucose intolerance as well as sleep disorders, together promoting obesity [29].

Night shift work in combination with overweight and obesity often lead to an increased waist circumference, so called abdominal obesity [22]. Waist circumference today is considered a good risk marker for chronic diseases such as high blood pressure, type 2 diabetes and cardiovascular disease [30]. Thus, measuring waist circumference is important when assessing the nutritional status of patients who perform night shifts. In the present study we also found a significant relationship between CD and body fat mass as well as visceral fat percentage. We found that the percentages of body fat mass and visceral fat are significantly higher in night shift workers. These findings are similar to those of the work of Son 2015 [31] and Sugiura 2020 [32], who also found an association between these parameters and inadequate physical activity as well as an increased risk of chronic diseases such as atherosclerosis.

#### 4.3. Sleep quality does not mediate deleterious effects of cronodisruption

Surprisingly, our findings suggest that sleep quality does not contribute to the association between CD and MS or between CD and BC. Our results contrast previous studies in the general population that have suggested that without taking night shift work into



consideration, sleep habits are associated with overweight and obesity [29]. These contradictions may be explained by differences in the duration of exposure to night shift work [11,33,34], unbalance of gender distribution among the comparison groups, or differences related to dietary habits, such as breakfast frequency 2019 [35]. Even though alterations of the metabolism can result from alterations in the sleep habits, napping can act as a protective factor for both weight gain and arterial hypertension [36,37]. Thus, it will be interesting to study the impact of napping on other chronic conditions such as MS.

#### 4.4. Strengths, limitations and methodological considerations

Given that the current study is the first of its kind in Ecuador, it has the potential to inspire further studies of MS and associated risk factors in the surrounding South American territories. Future studies should take into account various limitations we could not overcome. For example, we defined people exposed to CD as people who work in night shifts for at least 8 h in the last six months (see Methods). We did not measure the number of consecutive night shifts, exact duration of shifts or napping habits of night shift workers, which could have an impact on our conclusions [36–38]. Moreover, a more objective way to measure sleep quality could also be implemented. Thus, there are several improvements that could be implemented in future studies to control for potential caveats of the present study.

#### 4.5. General implications and recommendations

More studies should be performed to explain the current findings in a more comprehensive manner. Further research should focus on investigating the influence of genetic, physiological, environmental as well as life-style factors on shift work, MS, BC and effects of sleep habits. Together with a more objective way to measure sleep quality could also be implemented. The findings of this study may yield important implications for public health policies. For instance, people working in night shifts may want to consider to adjust their life style and/or to change their job after having lived in a reversed circadian rhythm for a longer period of time.

## 5. Conclusions

Our findings suggest that CD increases the odds of MS and influences BC in that, it increases the percentage of both fat mass and visceral fat. Interestingly, there was not sufficient evidence indicating that sleep quality mediates the deleterious impact of CD on health.

### Declaration of competing interest

The authors declare no conflict of interest.

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## Appendix A. Supplementary data

Supplementary data to this article can be found online at <https://doi.org/10.1016/j.dsx.2021.01.017>.

### Limitation of liability

The authors declare that all points of view expressed in this work are their entire responsibility and not the institution in which they work, or the source of funding.

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