

THE RELATIONSHIP BETWEEN WORKING SCHEDULE PATTERNS AND THE MARKERS OF THE METABOLIC SYNDROME: COMPARISON OF SHIFT WORKERS WITH DAY WORKERS

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Abstract

Objectives: This study examined the effect of shift work on developing the metabolic syndrome by comparing groups of exposed and unexposed Iranian drivers. **Methods:** We considered as night-shift drivers those drivers whose shifts included at least 15 h per week between 9:00 p.m. and 7:00 a.m. Daytime drivers were defined as drivers working regularly without shift work. 3039 shift work drivers were selected. These were matched with non-shift workers. The differences in baseline characteristics and the prevalence of the components of the metabolic syndrome were assessed with Student's t test, and chi-square tests. **Results:** We found central adiposity in 52.0% of the shift workers versus 42.6% of the day workers ($p < 0.0001$). The hypertension component was not significantly related to shift work ($p > 0.05$); but there were significant differences as regards other components of the metabolic syndrome ($p < 0.0001$). Among the shift workers, the odds ratios of the increased FBS, low HDL-C, higher TG levels, as well as higher waist circumference were 1.992 (95% CI: 1.697–2.337), 1.973 (95% CI: 1.759–2.213), 1.692 (95% CI: 1.527–1.874), and 1.460 (95% CI: 1.320–1.616), respectively. The metabolic syndrome was more common among the shift workers (OR = 1.495; 95% CI: 1.349–1.657). **Conclusion:** In evaluating such results, further consideration is needed to find pathophysiological clarification; in turn, stress linked to shift work must be considered to likely have had a relevant influence on the outcome. In our opinion, shift work acts as an occupational factor for the metabolic syndrome.

Key words:

Shift work, Insulin resistance, Metabolic syndrome, Abdominal obesity, Circadian clock

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INTRODUCTION

Human natural body rhythms are called circadian rhythms which are regulated by a “circadian clock”; located in the hypothalamus. This biological clock is synchronized by receiving the photic information from light-sensitive ganglion cells in the retina, thereby entraining individuals’ physiology and behavior to the external day–night cycle [1,2]. Nearly all of the biological processes including the sleep-wake cycles, body temperature, energy metabolism, cell cycle and hormone secretion have a circadian rhythm and are controlled by this circadian clock [2–5].

Shift work disrupts the clock’s function and is linked to circadian and metabolic consequences such as altered plasma lipid metabolism and adiposity [6].

A disruption of the circadian rhythm plays a key role in the pathophysiology of numerous diseases. It has become obvious that even disturbances of single aspects of these circadian rhythms may be associated with major effects [7–9]. It has been suggested that work/rest schedules resulting from shift work impair the physiological adaptation in the majority of night shift workers [1]. The sleep disorders, jet-lag syndrome, and increased proneness to work-related injuries are known short-term manifestation of shift work. In the long run, medical problems may manifest as increased risks for gastrointestinal, psychoneurotic, and cardiovascular diseases [10]. Shift work-related abnormalities in circulating lipids, central obesity, and hypertension have been repeatedly reported, but evidence of elevated blood pressure or metabolic disorders in a comparison between shift workers with day workers has not been convincingly demonstrated [6,11–15]. The metabolic syndrome (MetS) or the insulin resistance syndrome constitutes a clustering of several interrelated abnormalities that increase the risk for cardiovascular events and progression to diabetes mellitus [16,17]. On the other hand, the risk for a cardiovascular disease among shift workers is well documented. Several population studies indicate a relationship between shift work and morbidity from cardiovascular diseases [18].

Shift work may trigger the effect of lifestyle-related risk factors on coronary heart disease [19]. The present study was designed to determine the role of shift work impact on development of MetS, by comparing groups of exposed and unexposed Iranian professional drivers.

METHODS

Study population

Iranian health surveys of professional drivers of commercial motor vehicles have looked to improve their occupational health. From 2006 to 2010, more than 22 000 long-distance motor vehicle drivers were referred to our clinic. All of them reside in the West Azerbaijan Province of Iran. In this community-based study, a total of 6078 participants aged 20–60 years were included. The study protocol was approved by the Medical Ethics Committee of the Urmia University of Medical Sciences, and each participant signed a consent form.

Sampling method

For the purpose of the present study, we considered as night-shift drivers those drivers working on night or rotating shifts whose shifts included at least 15 h per week between 9:00 p.m. and 7:00 a.m. The daytime drivers were defined as drivers regularly working between 7:00 a.m. and 7:00 p.m. whose work was not shift-based. At baseline, a total of 4500 male shift work drivers aged 20–60 years were randomly selected among 22 453 individuals. We used simple random sampling because the population was uniform and had similar characteristics in all cases, e.g. occupation and the socioeconomic level.

Then, we excluded the drivers:

- who had worked as shift workers for less than two years,
- whose driving schedule included less than 44 h/week,
- whose shift work duration included less than 15 h/week from 9.00 p.m. to 7.00 a.m.

Finally, 3039 shift work drivers were selected. Thereafter, 3039 non-shift-work drivers were enrolled as a comparable group, who were matched with shift workers based on age, past smoking habit (pack-year history of smoking), period of employment as a driver, average of weekly driving in a year, and the type of the motor vehicle. Both groups were from similar socioeconomic backgrounds. The history of cigarette smoking was obtained from a questionnaire completed by each subject. The extent of smoking was quantified in pack-years, with 1 pack-years equivalent to 20 cigarettes per day for 1 year. Prior to the recruitment, all drivers had been informed about the aims of the health survey and the study. We excluded anyone who failed to provide us with an informed consent to participate in this study and applied the same inclusion and exclusion criteria to both groups.

Definition of MetS

Different organizations have proposed various criteria for the classification of MetS, and the National Cholesterol Education Program – Adult Treatment Panel III (NCEP-ATP III) as well as the International Diabetes Federation (IDF) criteria are used for the purpose of estimating the magnitude of MetS in the general population more widely than others. The IDF, in 2005, proposed a “consensus” definition of MetS in which waist circumference cut-off thresholds were made based on ethnicity specifics. All subjects were screened for MetS by using the IDF criteria that included the presence of central adiposity on the basis of waist circumference ≥ 94 cm plus two or more of the four following factors:

- elevated concentration of triglycerides (TG): ≥ 150 mg/dl (1.7 mmol/l),
- reduced concentration of HDL cholesterol: <40 mg/dl (1.03 mmol/l),
- elevated blood pressure: systolic blood pressure (SBP) ≥ 130 mmHg or diastolic blood pressure (DBP) ≥ 85 mmHg,

- elevated fasting plasma glucose (FBS) concentration ≥ 100 mg/dl (5.6 mmol/l) [20].

Medical history and clinical examination

According to the national protocol, demographic data including: age, shift work, smoking habits, duration of driving experience as the main job, type of motor vehicle, and the socioeconomic status of workers were collected using a standardized questionnaire. Waist circumference was measured at the level of the umbilicus in the standing position of the subjects. The subjects' blood pressure was measured after a five-minute rest, using a standard mercury sphygmomanometer with a suitable cuff calibrated by the Iranian Institute of Standards and Industrial Researches. Two consecutive diastolic and systolic blood pressures were recorded on the right arm. If the two-measurement difference exceeded 5%, blood pressure was measured for the third time. There were at least 30-second intervals between the measurements; the average of the two closest readings was considered as the subject's blood pressure.

Serum lipid and glucose analysis

Baseline blood samples were drawn into vacutainer tubes between 7:00 a.m. and 9:00 a.m. All the participants had fasted for ≥ 10 h. FBS and lipids were quantified at the time of specimen collection in the central laboratory of UMSU, a reference laboratory that had served Urmia residents for decades. Blood glucose and lipids, including TG and HDL cholesterol, were analyzed using a BT-3000 auto-analyzer (Biotecnica, Rome, Italy). The HDL-C and TG were assayed using enzymatic tests with commercially available kits (Pars Azmoon Inc., Iran). All samples were analyzed when the daily internal quality controls matched the acceptable criteria.

Statistical analysis

The characteristics of each group were summarized by means, standard deviations and percentages, subdivided

into four categories by age, and three body mass index (BMI = weight/height²) categories. The differences in baseline characteristics and the prevalence of components of MetS between the exposed and the non-exposed groups were assessed with Student's t test, and chi-square tests. A potential difference was considered significant on the basis of p, when it was lower than 0.05.

RESULTS

Among the components of Mets, the mean FBS, TG, and waist circumference in the group of shift workers were higher than in the group of day workers. These variables

differed significantly between the two groups. The shift workers group had significantly lower HDL. BMI differed significantly between the two groups, but the mean systolic and also diastolic BPs showed no significant differences between the two groups (Table 1). The frequency of overweight (BMI: 25–29.9) was the same in shift workers and day workers [1275 (42.0%) vs. 1279 (42.1%)]. In shift workers and day workers, respectively, obesity (BMI ≥ 30) was observed in 647 (21.3%) and 395 (13.0%) participants. The prevalence of the metabolic syndrome components in both groups is reported in Table 2. Central adiposity was the most common metabolic abnormality in both groups. The chi-square test revealed the central

Table 1. Baseline characteristics of the study population according to shift work

| Variable | Shift workers | | Day workers | | P |
|--------------------------|---------------|----------|-------------|----------|------------|
| | M | SD | M | SD | |
| Systolic BP (mmHg) | 123.71 | 14.729 | 123.57 | 14.428 | NS (0.721) |
| Diastolic BP (mmHg) | 77.4788 | 10.34743 | 77.7680 | 10.15777 | NS (0.272) |
| TG (mg/dl) | 182.88 | 143.992 | 152.13 | 101.456 | < 0.001 |
| HDL (mg/dl) | 44.13 | 9.895 | 46.26 | 9.279 | < 0.001 |
| FBS (mg/dl) | 91.58 | 21.084 | 88.90 | 15.533 | < 0.001 |
| Waist circumference (cm) | 93.6018 | 11.94843 | 90.5747 | 10.75563 | < 0.001 |
| BMI | 26.6760 | 4.09526 | 25.7145 | 3.76029 | < 0.001 |

BP – blood pressure, TG – triglyceride, HDL – high density lipoprotein, FBS – fasting blood sugar, BMI – Body Mass Index, M – mean, SD – standard deviation, NS – not significant.

Table 2. Prevalence of the components of the metabolic syndrome (MetS)

| Variable | Shift workers n (%) | Day workers n (%) | p | Odds ratio | 95% CI |
|---|------------------------|----------------------|------------|------------|-------------|
| FBS ≥ 100 mg/dl | 479 (15.8) | 261 (8.6) | < 0.0001 | 1.992 | 1.697–2.337 |
| HDL < 40 mg/dl | 1 052 (34.6) | 643 (21.2) | < 0.0001 | 1.973 | 1.759–2.213 |
| TG ≥ 150 mg/dl | 1 499 (49.3) | 1 110 (36.5) | < 0.0001 | 1.692 | 1.527–1.874 |
| Waist circumference ≥ 94 cm | 1 580 (52.0) | 1 294 (42.6) | < 0.0001 | 1.460 | 1.320–1.616 |
| Systolic BP ≥ 130 or diastolic BP ≥ 85(mmHg) | 1 089 (35.8) | 1 138 (37.4) | NS (0.101) | 0.933 | 0.840–1.036 |
| MetS | 1 384 (45.5) | 1 090 (35.9) | < 0.0001 | 1.495 | 1.349–1.657 |

Abbreviations as in Table 1.

adiposity in 52.0% of the shift workers versus 42.6% of the day workers ($p < 0.0001$). The hypertension component was not significantly related to shift work ($p > 0.05$); whereas there were significantly different frequencies of other components of MetS as well as BMI which are not part of the metabolic syndrome ($p < 0.0001$). Among the shift workers, the odds ratio of the increased FBS, low HDL-C, higher TG levels, as well as higher waist circumference were 1.992 (95% CI: 1.697–2.337), 1.973 (95% CI: 1.759–2.213), 1.692 (95% CI: 1.527–1.874) and 1.460 (95% CI: 1.320–1.616), respectively. Among the shift workers, MetS was more common than in the group of the day workers (odds ratio (OR): 1.495; 95% CI: 1.349–1.657).

In the shift workers and day workers, respectively, MetS increased from 25.8% and 24.5% within the 20–29 year-old group to 62.4% and 46.8% in the subjects of more than 50 years of age, as shown in Table 3.

The prevalence of two or more abnormalities of MetS without the obesity component was higher among the shift workers, whereas having one abnormality was manifested in the youngest age group of the day workers (Table 4).

Shift work increases the prevalence of MetS (OR = 1.495; 95% CI: 1.349–1.657). There was a significant age-related increase in the prevalence of MetS in both the shift and day workers groups. Our findings showed that it was more prevalent among the shift than day workers in all age groups.

DISCUSSION

After adjusting the results for objective confounders, the present study showed that the shift work pattern was associated with a higher prevalence of MetS. Our findings revealed that some components included in MetS, such as higher waist circumference, increased fasting blood glucose, and higher triglycerides blood levels as well as low HDL-C, were more common among the shift drivers in comparison to the day workers. On the other hand, we found no significant association concerning the blood pressure component between the shift workers and day ones.

The work-related factors of professional driving may affect the lipid metabolism and obesity due to various

Table 3. Prevalence of the metabolic syndrome (MetS) stratified by age groups and work category

| Working category | MetS | Age groups (years) / Year | | | | total |
|------------------|------|---------------------------|------------|------------|------------|--------------|
| | | 20–29 | 30–39 | 40–49 | > 50 | |
| Shift workers | yes | 160 (25.8) | 415 (40.4) | 481 (55.5) | 328 (62.4) | 1 384 (45.5) |
| | no | 461 (74.2) | 611 (59.6) | 385 (44.5) | 198 (37.6) | 1 655 (54.5) |
| Day workers | yes | 152 (24.5) | 369 (36.0) | 323 (37.3) | 246 (46.8) | 1 090 (35.9) |
| | no | 469 (75.5) | 657 (64.0) | 543 (62.7) | 280 (53.2) | 1 949 (64.1) |

Table 4. Prevalence of abnormalities of the metabolic syndrome (MetS), except the obesity component, stratified by work category

| Work category | | Frequency of components | | | | | total |
|---------------|--------------|-------------------------|--------------|------------|----------|---------------|-------|
| | | no | 1.00 | 2.00 | 3.00 | 4.00 | |
| Shift workers | 707 (23.3) | 1 039 (34.2) | 857 (28.2) | 378 (12.4) | 58 (1.9) | 3 039 (100.0) | |
| Day workers | 672 (22.1) | 1 626 (53.5) | 712 (23.4) | 14 (0.5) | 15 (0.5) | 3 039 (100.0) | |
| Total | 1 379 (22.7) | 2 665 (43.8) | 1 569 (25.8) | 392 (6.4) | 73 (1.2) | 6 078 (100.0) | |

harmful stressors including: the socioeconomic status, job and position, continuous tension caused by driving, lack of exercise, changes experienced in life due to an irregular work schedule [21–23]. Our findings are similar to those of other studies [24–26] that reported a higher tendency towards overweight or central obesity in the shift workers than in the day workers.

Increased blood pressure as the key component of MetS is the most widely studied factor in relation to the syndrome [27]. The relationship between elevated blood pressure and shift work is controversial. Although some studies have identified a trend towards higher blood pressure among shift workers [14], the present study and the results of other studies [26,28,29] showed no statistically significant differences in both the mean systolic and diastolic blood pressures.

In some studies, there was no difference in the triglyceride levels between the shift and day workers [12,26], whereas several population studies have shown that shift work could lead to a high prevalence of the elevated triglyceride levels concentrations [13,6]. It has been suggested that shift work is associated with an increase in the triglyceride levels independent of the dietary intake [30]. In the present study, the same was true for higher triglyceride levels among the shift workers. Karlsson et al. found a significant association between shift work and obesity, and also the abnormalities in circulating lipids such as low HDL-cholesterol and high triglyceride levels, indicating an association between shift work and the metabolic syndrome. Based on their findings, the risk of low HDL-cholesterol was doubled among the shift workers [6].

Although the abnormalities in circulating lipids and abdominal obesity have been repeatedly reported in shift workers, there are few studies reporting impaired glucose metabolism in relation to shift work. In the present study, higher FBS in shift workers was consistent with a 14-year cohort study of 7104 male Japanese workers which

revealed that shift work was an independent risk factor for impaired glucose metabolism [31].

Although the primary cause of the weight gain leading to MetS is attributed to increased caloric intake and decreased energy expenditure, some studies point out that the overall food content or caloric intake are not related to shift work [32].

Epidemiological studies have documented that abdominal obesity and its metabolic comorbidities are the consequence of a chronic maladaptation to environmental stress factors [33], which is also in agreement with some epidemiological studies showing an association of MetS with the job stress [34,35]. The hallmark of the stress response is increased cortisol secretion by the hyperactivation of the hypothalamic-pituitary-adrenocortical (HPA) axis [36]; which can cause insulin sensitivity to decrease in the muscles [37], inhibit glucose uptake by the peripheral tissues and also lead to an increase in both lipolysis and gluconeogenesis [38].

Although we cannot define this condition precisely as a direct effect of shift work, it should be noted that visceral obesity plays the key role in the development of insulin resistance [39], and insulin levels are associated with higher BMI and also higher visceral obesity which alters glucose metabolism mediated by decreasing insulin sensitivity [24]. Shift work increases obesity [6], so the difference in the blood glucose levels between the shift workers and the control group may be indirectly caused by shift work. In addition, the measurement of BMI could be useful for identifying persons at increased risk of developing MetS among the shift workers. The same may be true for the disruption of the circadian rhythm which may “per se” play the main role in inducing metabolic disturbances of MetS [11,40]. It is worth noting that some investigators have reported that metabolic changes have been observed to occur independently of poor diet and physical inactivity. Branth et al. concluded that despite regular exercise and a proper diet, subjects under

prolonged stress developed metabolic alterations including distinctive central obesity, biochemical changes and slight hypertension towards MetS [41]. As an opposite reason as in the earlier studies [6,12,24,41], it seems possible that shift work is responsible for both the elevated TG levels and higher FBS. In turn; stress linked to shift work causes insulin sensitivity to decrease and impaired FBS.

In the present study, we did not examine dietary habits and daily exercise with objective methods; so, no clear-cut conclusion could be drawn with regard to this aspect. Professional driving is associated with substantial changes in lifestyle habits. Physical inactivity is common in Iranian long-distance drivers and, also, an interview conducted with some subjects by a dietitian clearly showed that the only option they have is to have their meals in restaurants. Both shift and day workers have similar limited options as regards their meals. Most restaurants use high saturated fat for daily cooking and the usual meals consumed in these restaurants are poor in fibers (vegetables and fruits), rich in animal proteins (beef, sheep, veal), foods with high glycemic index (rice, white bread), and saturated fat (cheese, eggs) that are well known to induce an increase of the body fat in working individuals [42].

The present study supports the effect of shift work on MetS in line with those from other authors. In evaluating such results, further consideration is needed to find pathophysiological clarifications; in turn, stress linked to shift work, must be considered as having a relevant influence on MetS.

CONCLUSION

In our opinion, shift work acts as an occupational factor for MetS. All professional drivers of commercial motor vehicles, particularly shift workers, should be carefully evaluated by occupational physicians for MetS in order to promote preventive and corrective actions.

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