Association between rotating night shift work and metabolic syndrome in Korean workers: differences between 8-hour and 12-hour rotating shift work

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Abstract: This study aimed to analyze the association between the shift work schedule and metabolic syndrome (MetS). This is a retrospective longitudinal study based on the 2015 health checkup data of 2,090 workers evaluated for MetS in 2010 at a general hospital in Korea. The participants were divided according to their shift work schedule into daytime, three-shift (8-h rotation), and twoshift (12-h rotation) workers. The index that indicates the association between the shift work schedule and MetS is the odds ratio (OR) calculated using multivariate logistic regression. The analysis for the entire group of workers indicated that there was positive association between two-shift rotation and MetS (OR=1.58, 95% confidence interval [CI]: 1.09, 2.29). In the analysis of rotating nightshift workers, the years of rotating night shifts, frequency of night-shift work, and sleep disturbance were added to the confounding variables, and two-shift work remained positively associated with MetS (OR=1.72, 95% CI: 1.10, 2.70). The risk of MetS differs based on the shift work schedules they engage in. Hence, structural changes to the shift work schedules are required to prevent MetS in night-shift workers.

Key words: Metabolic syndrome, Rotating night shift work, Retrospective longitudinal study, Three-shift rotation, Two-shift rotation

Introduction

The 24-hour-a-day production schedule of the manufacturing industry and changes to the human use of services have made night-shift work mandatory. In Europe, 19% of the entire workforce engages in night-shift work¹). In Korea, according to the Ministry of Employment and Labor's 2011 research that sampled 3,413 firms, 15.2% of the entire workforce was engaged in night-shift work²). Rotation shifts that include night shifts are known to predispose to obesity^{3, 4}), high blood pressure^{5, 6}), dyslipid-

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emia⁷⁾, and diabetes^{8, 9)}. From 2014, the Korean government made monitoring of night-shift work exposure and the health of night-shift workers mandatory.

Metabolic syndrome (MetS) is defined as a cluster of risk factors for diabetes and cardiovascular diseases (CVDs). The prevention of MetS is believed to decrease the prevalence of diabetes and CVDs, and as such, MetS prevention is an important health focus worldwide¹⁰). While many studies report that rotating night-shift work is a risk factor for MetS, this finding is not consistent^{11–16}). Since the scheduling of night-shift work determines the frequency and extent to which the worker is exposed to night-shift work, it is an important workplace factor for evaluating the risk of MetS. However, very few studies have classified night-shift work by rotation in order to examine the

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Fig. 1. Selection of study participants.

association between rotating night-shift work and MetS. Thus, the aim of this study was to evaluate the association between rotating shift schedule and MetS.

Methods

Study design and population

This retrospective longitudinal study utilized worker health checkup data obtained on a per-workplace basis from a tertiary hospital located in Ansan City, Korea. Thirty-one workplaces had their workers evaluated for MetS at the hospital in 2010. They were all manufacturing companies located in the Ansan-Siheung industrial complex and with rotating shift schedule. These workplaces also conducted health checkups for their workers at the same hospital in 2015. A total of 3,051 workers aged between 20 and 59 yr who were assessed for MetS in 2010 were selected. Of these, 2,090 were included in this study after excluding workers with the following conditions: employees who did not undergo health checkups in 2015 at this hospital (n=510); those who were diagnosed with MetS in 2010 (n=269); those who had a change of the shift work schedule between 2010 and 2015 (n=71); those who had some missing test items necessary for diagnosis of MetS in 2015 (n=43); or those who lacked data on confounding variables for analysis (n=68) (Fig. 1).

The 2010 and 2015 data of occupational factors such as engagement in night-shift work, shift work schedule, and years of rotating night shifts were provided by the managers of the employee health affairs of each company. The workers who engaged in night-shift work are required to fill in the questionnaire for night-shift work during their health checkups. This questionnaire contained questions such as shift work schedule, years of rotating night shifts, and frequency of night-shift work. The 2015 data of this questionnaire were used as supplemental data in this study. The physician rechecked all examinees regarding their engagement in night-shift work, shift work schedule, type of job, years of rotating night shifts, frequency of nightshift work, and change of shift work schedule between 2010 and 2015 during the medical examination. Health checkups were carried out in the morning at the respective workplaces. The medical staff instructed the workers to fast for at least 8 h prior to the health checkup. This study was approved by the Institutional Review Board of the Catholic University of Korea, and the requirement for informed consent was waived since data were collected retrospectively without any participant-identifying information.

Assessment of rotating night-shift work

This study defined night-shift work and night-shift workers as per the regulations of the Ministry of Employment and Labor of Korea¹⁷⁾.

- Night-shift work: Work performed between 10 p.m. and 6 a.m.
- Night-shift worker: A worker that satisfies either of the following two criteria:
 - 1. Engaged in night-shift work for over 60 h per month over 6 months.
 - 2. Carried out an average of four or more shifts of work between 10 p.m. and 6 a.m. continuously, over the period of 6 months.

Night-shift work was comprised of either three 8-h rotating shifts or two 12-h rotating shifts. The 8-h rotating shift was made up of a morning, an evening, and a night shift. The 12-h rotating shift consisted of a day shift and a night shift. Of the 31 workplaces, 8 operated an 8-h rotating shift pattern; 21, a 12-h pattern; 2, both shift patterns.

Assessment of MetS

The diagnosis of MetS in this study was based on the modified NCEP ATP III criteria¹⁸). A worker was diagnosed with MetS if they satisfied three or more out of the five following criteria: 1) elevated blood pressure $\geq 130/85$

mmHg or receiving treatment for hypertension; 2) elevated fasting plasma glucose level ≥ 100 mg/dl or receiving treatment for elevated blood glucose; 3) high serum level of triglyceride ≥ 150 mg/dl or receiving treatment for dyslipidemia; 4) low serum level of high-density lipoproteins (HDL) cholesterol: men <40 mg/dl, women <50 mg/dl or receiving treatment for dyslipidemia; 5) waist circumference: men ≥ 90 cm, women ≥ 80 cm. The same equipment and methods were used for measuring MetS components in 2010 and in 2015. Waist circumference and blood pressure were measured by nurses, and fasting glucose, triglyceride, and HDL cholesterol were measured using Modular P800 (Roche, Swiss).

Assessment of confounding variables

Smoking status, alcohol consumption, and the physical activity of workers were evaluated using medical history taking and self-administrated questionnaires in 2015. Smoking status was classified into current smokers and non-smokers. Current smoker is defined as a person who has smoked 100 cigarettes in his or her lifetime and who currently smokes cigarettes. Using the criteria of the National Institute on Alcohol Abuse and Alcoholism, alcohol consumption was divided into three groups: none, lowrisk drinking, and high-risk drinking. We defined highrisk drinking as ≥ 5 glasses/d or ≥ 15 glasses/wk for men and ≥ 4 glasses/d or ≥ 8 glasses/wk of alcohol for women. We defined one glass as 50 ml of soju (distilled liquor in Korea). Physical activity levels were measured using the Korean version of the International Physical Activity Questionnaire (IPAQ), short form¹⁹⁾. According to IPAQ, physical activity can be categorized into three levels: high, moderate, and low²⁰⁾. In this study, if the physical activity of workers were moderate or above (≥600 MET-min/ wk), this was classified as sufficient physical activity. The size of the workplace was classified into large (\geq 300 workers) and medium (50-299 workers). There were no small workplaces with <50 workers among the workplaces selected for the study. The type of job performed by the workers in this study was either white or blue collar. White-collar work included office, research, and management roles; blue-collar work included manufacturing, operation and maintenance, and transportation and logistics roles. The data of years of rotating night shifts and frequency of night-shift work were collected in the form of continuous variables. The frequency of night-shift work is evaluated as the average number of night-shift per month over a period of 6 months. Evaluation of sleep disturbance was carried out in night-shift workers using the Insomnia

Severity Index $(ISI)^{21}$. The Korean version of the ISI is reported to be a reliable and valid instrument for assessing insomnia severity in Koreans²²⁾. The total score is interpreted as follows: 0–7, no clinically significant insomnia; 8–14, subthreshold insomnia; 15–28, clinical insomnia.

Statistical analysis

For the statistical analysis, the study participants were divided into daytime, three-shift (8-h rotation), and twoshift (12-h rotation) workers. We evaluated the differences among the groups using chi-square tests for categorical variables, and we also conducted post hoc tests after chisquare tests using Dunnett's test. The t-test was used for continuous variables between two- and three-shift rotations. Confounding variables were adjusted using multivariate logistic regression. Variables shown to have a statistically significant relationship with MetS in the univariate logistic regression ($p \le 0.2$) were included in the multivariate logistic regression model as confounding variables. The results after adjusting for confounding variables are shown as odds ratio (OR) for three- and two-shift rotation, with the daytime as the reference group. First, variables were added in phases for different models for all the workers: model 1, adjusted for age and sex; model 2, additionally adjusted for smoking status (current smoker/non-smoker), alcohol consumption (no/social/heavy), and physical activity (sufficient/insufficient); model 3, additionally adjusted for workplace size (≥300 workers/50-299 workers) and type of job (white-collar/blue-collar). Additional analysis using multivariate logistic regression was performed for the rotating night-shift workers alone. For rotating night-shift workers, the three-shift rotation was used as the reference group, and the OR for the two-shift rotation was calculated. Moreover, additional variables of years of night-shift rotating, frequency of night-shift work, and sleep disturbance were added, which were not relevant to daytime workers: model 1, adjusted for age and sex; model 2, additionally adjusted for smoking status, alcohol consumption, and physical activity; model 3, additionally adjusted for workplace size and type of job, years of rotating night shifts (continuous), frequency of night-shift work (continuous), and sleep disturbance (no/subclinical/ clinical). Of 3,051 workers, 510 who were evaluated for MetS in 2010 were lost to follow-up in the 2015 health checkup. For these 510 workers, only sex, age, and MetS prevalence rate of 2010 were confirmed. With regard to the MetS prevalence rate in 2010, sex, and age distribution, 510 workers lost to follow-up were compared with the 2,541 workers who received health checkups at the hospital in 2015 using chi-square test. SPSS 18 (IBM SPSS Inc., Chicago, IL, USA) was utilized for the statistical analyses.

Results

The study participants (n=2.090) were divided into three groups depending on their shift work schedule: daytime, three-shift rotation, and two-shift rotation. Among the participants, 907 worked daytimes, 538 worked threeshift rotation, and 645 worked two-shift rotation. The proportion of female and white-collar workers was higher in the daytime group than in the other two groups. The twoshift rotation group had the lowest proportion of workers who work at large-size workplaces among the three groups. Otherwise, the smoking and drinking rates of the two-shift rotation group were higher than the other two groups. The three-shift rotation group was older than the other two groups and had the highest percentage of workers in large-size workplaces. The years of rotating night shifts and the frequency of night-shift work were longer and higher, respectively, in the three-shift rotation group than in the two-shift rotation group (Table 1).

In multivariate logistic regression analysis conducted with the three groups, two-shift rotation was positively associated with MetS in all models: model 1, OR=1.53, 95% confidence interval [CI]: 1.13, 2.09; model 2, OR=1.51, 95% CI: 1.11, 2.06; model 3, OR=1.58, 95% CI: 1.09, 2.29. However, there was no association between three-shift rotation and MetS in any of the models (Table 2).

The multivariate logistic regression analysis including the rotating night-shift workers alone, where the years of rotating night shifts, the frequency of the night-shift work, and sleep disturbance were added as confounding variables, continued to indicate that two-shift rotation was associated with MetS (OR=1.72, 95% CI: 1.10, 2.70) (Table 3).

The results of the comparison between workers lost to follow-up and workers who received health check in 2015 showed a significant difference only in age distribution. Among the workers lost to follow-up, the percentage of workers in their 20 s was high (Table 4).

Discussion

A leading mechanism that explains the association between night-shift work and MetS is circadian misalignment. The suprachiasmatic nucleus of the anterior hypothalamus controls the endogenous circadian rhythm, which

| Variables | Daytime (n=907) | Three-shift rotation $(n=538)$ | Two-shift rotation ($n=645$) | p-value ^a |
|---|-----------------|--------------------------------|--------------------------------|----------------------|
| Sex | | | | < 0.001*** |
| Male | 775 (85.4) | 495 (92.0) | 602 (93.3) | |
| Female | 132 (14.6) | 43 (8.0) | 43 (6.7) | |
| Age (years) | | | | < 0.001*** |
| 25-34 | 96 (10.6) | 28 (5.2) | 38 (5.9) | |
| 35-44 | 399 (44.0) | 187 (34.8) | 334 (51.8) | |
| 45-54 | 292 (32.2) | 257 (47.8) | 212 (32.9) | |
| 55-64 | 120 (13.2) | 66 (12.3) | 61 (9.5) | |
| Smoking status | | | | < 0.001*** |
| Non-smoker | 605 (66.7) | 334 (62.1) | 352 (54.6) | |
| Current smoker | 302 (33.3) | 204 (37.9) | 293 (45.4) | |
| Alcohol consumption | | | | 0.015* |
| None | 283 (31.2) | 189 (35.1) | 174 (27.0) | |
| Low-risk drinking | 207 (22.8) | 98 (18.2) | 139 (21.6) | |
| High-risk drinking | 417 (46.0) | 251 (46.7) | 332 (51.5) | |
| Physical activity | | | | 0.360 |
| Sufficient | 427 (47.1) | 271 (50.4) | 300 (46.5) | |
| Insufficient | 480 (52.9) | 267 (49.6) | 345 (53.5) | |
| Workplace size | | | | < 0.001*** |
| Medium(50-299 workers) | 321 (35.4) | 141 (26.2) | 278 (43.1) | |
| Large (≥300 workers) | 586 (64.6) | 397 (73.8) | 367 (56.9) | |
| Type of job | | | | < 0.001*** |
| White-collar | 539 (59.4) | 50 (9.3) | 30 (4.7) | |
| Blue-collar | 368 (40.6) | 488 (90.7) | 615 (95.3) | |
| Metabolic syndrome | | | | 0.009** |
| Yes | 93 (10.3) | 59 (11.0) | 98 (15.2) | |
| No | 814 (89.7) | 479 (89.0) | 547 (84.8) | |
| Sleep disturbance ^c | | | | 0.170 |
| No | | 317 (58.9) | 355 (55.0) | |
| Subthreshold insomnia | | 170 (31.7) | 208 (32.2) | |
| Clinical insomnia | | 51 (9.5) | 82 (12.7) | |
| Years of rotating night shifts (year) ^{bc} | | 15.8 ± 6.3 | 13.5 ± 5.9 | 0.02* |
| Frequency of night shift work ^{bc} | | 8.3 ± 0.7 | 10.3 ± 1.7 | < 0.001*** |

Table 1. Characteristics of the sample by shift work schedule

Values for categorical variables given as number (percentage); Values for continuous variables given as mean \pm SD.

p < 0.05, p < 0.01, p < 0.001

^a χ^2 test, categorical variables; *t*-test, continuous variables.

^b Continuous variables.

^c Variables for rotating night shift workers.

| Table 2. | Odds ratios of | f metabolic | syndrome | by the | shift work schedule ^a | |
|----------|----------------|-------------|----------|--------|----------------------------------|--|
|----------|----------------|-------------|----------|--------|----------------------------------|--|

| | Model 1 ^b | | | Model 2 ^c | | Model 3 ^d | |
|----------------------|----------------------|-------------------|------|----------------------|------|----------------------|--|
| | OR | 95% CI | OR | 95% CI | OR | 95% CI | |
| Daytime | 1 | Reference | 1 | Reference | 1 | Reference | |
| Three-shift rotation | 0.96 | (0.68 - 1.36) | 0.97 | (0.68 - 1.38) | 0.98 | (0.66 - 1.45) | |
| Two-shift rotation | 1.53 | $(1.13 - 2.09)^*$ | 1.51 | $(1.11 - 2.06)^*$ | 1.58 | $(1.09 - 2.29)^*$ | |

Abbrevations: OR, odds ratio; CI, confidential interval.

*p<0.05, **p<0.01, ***p<0.001.

^a ORs were calculated by multivariate logistic regression.

^b Model 1 was adjusted for age and sex.

^c Model 2 was additionally adjusted for smoking status (current smoker/non-smoker), alcohol consumption (no/social/heavy) and physical activity (sufficient/insufficient).

^d Model 3 was additionally adjusted for workplace size (\geq 300 workers/50–299 workers) and type of job (white/blue-collar).

| | Model 1 ^b | | | Model 2 ^c | | Model 3 ^d | |
|----------------------|----------------------|-------------------|------|----------------------|------|----------------------|--|
| | OR | 95% CI | OR | 95% CI | OR | 95% CI | |
| Three-shift rotation | 1 | Reference | 1 | Reference | 1 | Reference | |
| Two-shift rotation | 1.63 | $(1.15 - 2.33)^*$ | 1.61 | $(1.13 - 2.29)^*$ | 1.72 | $(1.10-2.70)^*$ | |

Table 3. Odds ratios of metabolic syndrome by shift work schedule among night shift worker^a

Abbrevations: OR, odds ratio; CI, confidential interval.

p < 0.05, p < 0.01, p < 0.01, p < 0.001.

^a ORs were calculated by multivariate logistic regression.

^b Model 1 was adjusted for age and sex.

^c Model 2 was additionally adjusted for smoking status (current smoker/non-smoker), alcohol consumption (no/social/heavy) and physical activity (sufficient/insufficient).

^d Model 3 was additionally adjusted for workplace size (\geq 300 workers/50–299 workers), type of job (white/blue-collar), the years of rotating night shifts (continuous), the frequency of night shift work (continuous), and sleep disturbance (no/subclinical/clinical).

 Table 4.
 Characteristics of the population who participated in the 2010 health checkups

| | Total (n=3,051) | Participated heal | | |
|--------------------|--------------------|----------------------------|----------------------|-----------------|
| Variables | | 2010 and 2015 (n=2,541) | 2010 only (n=510) | <i>p</i> -value |
| Gender | | | | 0.557 |
| Male | 2,770 (90.8) | 2,303 (90.6) | 467 (91.6) | |
| Female | 281 (9.2) | 238 (9.4) | 43 (8.4) | |
| Age (years) | | | | < 0.001*** |
| 20-29 | 336 (11.0) | 187 (7.4) | 149 (29.2) | |
| 30-39 | 1,286 (42.2) | 1,097 (43.2) | 189 (37.1) | |
| 40-49 | 1,060 (34.7) | 946 (37.2) | 114 (22.4) | |
| 50-59 | 369 (12.1) | 311 (12.2) | 58 (11.4) | |
| Metabolic Syndrome | | | | 0.209 |
| Yes | 324 (10.6) | 278 (10.9) | 46 (9.0) | |
| No | 2,727 (89.4) | 2,263 (89.1) | 464 (91.0) | |

Values for categorical variables given as number (percentage).

p-values calculated using χ^2 test.

p*<0.05, *p*<0.01, ****p*<0.001

affects the activity of a number of hormones such as the growth hormone, melatonin, cortisol, leptin, and ghrelin^{23, 24)}. As such, circadian misalignment between the endogenous circadian rhythm and the 24-h behavioral cycles caused by rotating night shifts disrupts the normal function of these hormones and negatively affects the maintenance of homeostasis of glucose and lipids^{24, 25)}. Several studies also report that circadian misalignment is common in shift workers, leading to increased blood glucose levels and blood pressure^{25, 26)}. Two-shift (12-h rotating shifts) workers have longer working hours in the night shift compared with three-shift (8-h rotating shift) workers. Moreover, two-shift work is composed of day and night shifts; thus, the frequency of night-shift work for workers are also greater compared to three-shift work, which includes morning, evening, and night shifts. Given such

characteristics, it is possible that circadian misalignment occurs at a higher severity and proportion in two-shift workers compared with three-shift workers. This study also showed that the frequency of night-shift work for twoshift workers was higher than that for three-shift workers. We have included the frequency of night-shift work as a confounding variable in multivariate analysis. However, even after adjusting for the frequency of night-shift work, the risk of MetS was still increased in two-shift workers. Circadian misalignment is strongly associated with sleep deprivation on night-shift worker²⁷⁾, and many studies suggest that sleep deprivation, comparatively more common for night-shift workers, is associated with MetS²⁸⁻³⁰⁾. We used ISI in evaluating sleep disturbance, which has proven validity and reliability as an index for evaluating responses to insomnia treatment and as a screening instrument during primary care³¹⁾. However, in this study, the three- and two-shift workers had no meaningful differences in sleep disturbance.

The results of smoking habit, alcohol consumption, and physical activity in this study were similar to those of other studies. Previous studies have reported that night-shift workers are more likely to smoke and drink alcohol^{32–34}, and many studies have shown that physical activity does not differ between daytime and night-shift workers^{35–37}, similar to the results of this study. We could not obtain the data on the income levels of the workers in this study. In general, larger workplaces have higher income levels compared with smaller ones. The 2015 Labor Study on Employment Types also reported higher incomes for workplaces with <300 workers³⁸. As such, with the multivariate analysis adjusting the effects of workplace size, the influence of income levels is expected to adjust accordingly.

We have only confirmed one existing study conducted in Japan that evaluated the risk of MetS based on shift rotation type³⁹⁾. However, contrary to this present study, the study in Japan reported that two-shift work was negatively associated with MetS (OR=0.77, 95% CI: 0.61, 0.98). We retrospectively used data from 2010 and could exclude workers who were diagnosed with MetS in 2010 in the analysis, but the Japanese study was cross-sectional. In addition, the present study followed the criteria of the modified NCEP ATP III for diagnosing MetS; however, the aforementioned study adhered to the standard Japanese criteria for diagnosing MetS. The standard Japanese criteria for the diagnosis of MetS, unlike the modified NCEP ATP III, have an essential component of waist circumference and have a higher fasting blood glucose standard (≥ 110 mg/dl). Thus, it was difficult to compare the results of that study with those of the present study. The results of another study that examined daytime and two-shift workers were similar to the results of this study^{14, 40}. Studies that examined daytime and three-shift workers show different results between researchers. A study conducted in Finland found that the risk of MetS in three-shift workers did not increase compared to that of daytime workers, as observed in this current study⁴¹; however, in a study conducted in France on 190 workers, three-shift work was positively associated with MetS¹³⁾. In this current study, 73.8% of the three-shift workers belonged to large workplaces (\geq 300 workers). We believe that the difference of workplace size may have led to the differences in the results of the two studies.

The strength of this present study is that, unlike other studies on night-shift work and MetS, it further classified rotating night-shift workers into three- and two-shift workers to confirm their respective associations with MetS. Moreover, in this study, the classification of night-shift and daytime workers was carried out by the manager of employee health affairs of each company. Additionally, we rechecked their engagement in night-shift work and their working histories. As such, there was a lower possibility of misclassification for night-shift. Another strength of this study can be found in the participants, who worked with the same shift work schedule at the same workplace from 2010 to 2015. If retired workers and workers who were temporarily involved in rotating night-shift work are included in the study, the changes in shift work schedule may affect the outcome.

This study has several limitations. First, some of the confounding variables included in the analysis are crosssectional data. Second, we could not include the data of the workers' education level as a confounding variable. Lastly, there is no information on the workers' eating habits. However, the results of studies on the differences of the calorie intake between daytime and night-shift workers differ from each other^{13, 42, 43)}. Results also differed in terms of meal frequency^{13, 44)}. This study was conducted with two- and three-shift workers as exposure groups who had rotating night-shift work for more than 5 yr, and two- and threeshift work are the most common types of rotating nightshift work. In addition, different types of manufacturing companies were included in this study. Therefore, it is possible to apply its results to rotating night-shift workers at manufacturing-related workplaces with ≥ 50 workers. However, caution must be exercised in applying the results of this study to workers in other professions as well as to workers in workplaces with < 50 workers.

In conclusion, the results of this study indicate that for night-shift workers, the risk of MetS would differ based on the shift work schedules they engage in. To prevent MetS in night-shift workers, structural changes to the shift work schedules are required, in addition to individual efforts on the part of workers such as lifestyle modifications.

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Conflict of Interest

None declared.

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