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# Relationship between chronic rhinosinusitis and risk of obstructive sleep apnea

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The relationship between obstructive sleep apnea (OSA) and chronic rhinosinusitis (CRS) has not yet been fully elucidated. Therefore, the objective of this study was to evaluate the connection between OSA risk and CRS by investigating associations between the STOP-Bang questionnaire and presence of CRS in a nationwide, population-based study. This is a cross-sectional study based on the Korean National Health and Nutrition Examination Survey (KNHANES). We evaluated 10,081 subjects who completed both the STOP-Bang and CRS-related questionnaires. Among the total subjects, 390 (3.9%) were CRS patients. The median STOP-Bang score was 3.0 [2.0; 4.0] in CRS patients, compared to 2.0 [1.0; 3.0] in subjects without CRS. In a low-risk group according to the STOP-Bang questionnaire, 3.1% of subjects were CRS patients. However, a gradual increase was observed among different risk groups. In the higher risk group, CRS patients accounted for 5.3% (P < 0.001). Among the four main symptoms of CRS (nasal obstruction, nasal discharge, facial pain/pressure, and decreased sense of smell), nasal obstruction (4.1 to 7.3%) and a decreased sense of smell (1.9 to 3.3%) increased with higher STOP-Bang scores. This study found that the proportion of patients with CRS was significantly higher in the group at a higher STOP-Bang score in the general population. Among symptoms of CRS, nasal obstruction and anosmia were found to be associated with an increased STOP-Bang score.

Keywords Sinusitis, Obstructive sleep apnea, Risk, Questionnaire, KNHANES

Obstructive sleep apnea (OSA) is a disorder in which multiple upper airways are intermittently blocked during sleep<sup>1,2</sup>. The estimated prevalence of moderate to severe OSA requiring treatment varies depending on age group and sex, ranging from 3 to 50%<sup>3,4</sup>. OSA is known to be an independent risk factor for cardiovascular and cerebrovascular diseases, both of which are associated with an individual's mortality<sup>5</sup>. Consequently, the importance of OSA diagnosis is steadily increasing.

Recent studies have suggested that chronic intermittent hypoxia due to OSA can activate inflammatory pathways and increase oxidative stress, as indicated by high serum levels of interleukin (IL)-1, IL-6, IL-8, and tumor necrosis factor (TNF)- $\alpha^{6-8}$ . The systemic inflammatory mechanisms of OSA are well-documented for many chronic diseases such as hypertension, diabetes mellitus (DM), and other metabolic syndromes. It has been suggested that OSA is also associated with chronic inflammatory diseases such as CRS<sup>9-11</sup>. Chronic rhinosinusitis (CRS) refers to the inflammation of the nose and paranasal sinuses. The role of the nose in OSA is already well known because people breathe through the nose during sleep<sup>12,13</sup>. Moreover, previous studies have reported that OSA and sleep disruption are highly prevalent in patients with CRS<sup>14,15</sup>. Currently, there is a lack of research assessing how the presence and symptoms of CRS in the general population are related to the STOP-Bang questionnaire.

The gold standard diagnostic tool for OSA is polysomnography<sup>16</sup>. However, due to limited access to sleep laboratories, OSA is underdiagnosed in the general population<sup>17</sup>. Therefore, the STOP-Bang questionnaire is widely used as an alternative screening test. The STOP-Bang questionnaire is a concise and easy-to-use screening tool for identifying patients at high risk of underlying moderate to severe OSA<sup>15,16</sup>. The STOP-Bang questionnaire was designed as a screening tool for OSA, comprising 4 self-reportable factors (STOP: snoring, tiredness, observed apnea, and high blood pressure) and four demographic variables (Bang: body mass index (BMI), age, neck circumference, and gender)<sup>18</sup>. A STOP-Bang score of 3 or higher demonstrates excellent sensitivity in

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discriminating mild, moderate, and severe OSA (sensitivity: 84%, 93%, and 100%, respectively in the original validation trial)<sup>18,19</sup>. The diagnostic accuracy of a STOP-Bang score of 3 or higher in detecting moderate to severe OSA was greater than 0.80<sup>18</sup>. The significance of STOP-Bang lies not only in its ability to measure the risk of OSA, but also in its role as a predictor of detrimental outcomes such as cardiovascular risk associated with OSA<sup>20</sup>.

The Korea National Health and Nutrition Examination Survey (KNHANES) is an annual survey conducted by Korea Disease Control and Prevention Agency to assess nationwide health and nutrition statistics<sup>21</sup>. This nationally representative cross-sectional survey annually includes approximately 10,000 individuals and gathers data on socioeconomic status, health-related behaviors, quality of life, clinical profiles relevant to non-communicable diseases, and other related factors<sup>22</sup>. The increasing clinical importance of OSA prompted the inclusion of the STOP-Bang questionnaire as a survey item for adults aged 40 and above starting in 2019 (The KNHANES VIII)<sup>21</sup>. Therefore, the aim of this study was to assess the relationship between OSA risk and CRS by analyzing correlations between the STOP-Bang questionnaire and CRS presence and symptoms in the nationwide study.

## Results

#### Relationship between presence of CRS and STOP-Bang questionnaire

The KNHANES survey period for the STOP-Bang and CRS-related questions was from 2019 to 2021. Out of the 22,559 individuals sampled during this period, 9,541 who did not complete the STOP-Bang questionnaire and an additional 2937 individuals who did not complete the CRS-related questionnaire were excluded. Therefore, a total of 10,081 subjects were included in this study. The final sample consisted of 390 (3.9%) individuals with CRS and 9691 individuals without CRS (Fig. 1). The median STOP-Bang score was 3.0 for patients with CRS (Mean ± standard deviation;  $2.8 \pm 1.6$ ) and 2.0 for those without CRS ( $2.5 \pm 1.5$ ) (Fig. 2). The power size was 0.963. CRS patients exhibited higher percentages of snoring (25.4%), tiredness (46.2%), observed apnea (11.3%), and thick neck circumference (41.8%) compared to subjects without CRS (17.3%, 27.0%, 7.8%, 35.2%, respectively). However, age was significantly lower in patients with CRS (median age, with CRS: 58.0, without CRS: 60.0; P < 0.001). Additionally, the percentage of individuals over the age of 50 was significantly higher in the CRS group. This contrasts with the known association of increasing age with a higher risk of OSA (Table 1). Cigarette smoking was significantly (49.2% of CRS patients and 39.6% of subjects without CRS; P < 0.001) more prevalent in

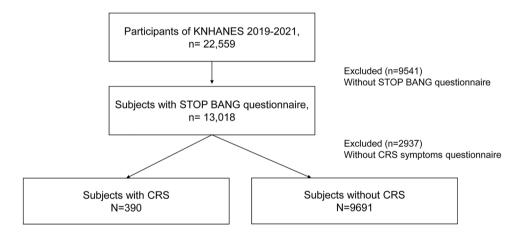


Fig. 1. Flow diagram of participant recruitment.

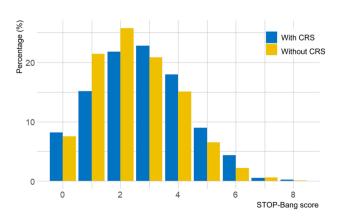


Fig. 2. Distribution of STOP-Bang score by chronic rhinosinusitis status.

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	Patients, by presence of CRS			
	with CRS (N = 390)	without CRS (N=9691)	P value	
STOP-Bang (0-8)	3.0 [2.0; 4.0]	2.0 [1.0; 3.0]	< 0.001	
STOP				
Snore	99 (25.4%)	1680 (17.3%)	< 0.001	
Tired	180 (46.2%)	2616 (27.0%)	< 0.001	
Observed	44 (11.3%)	754(7.8%)	0.016	
Pressure	129 (33.1%)	3392 (35.0%)	0.467	
Bang		1		
BMI	24.1 [21.6;26.3]	24.0 [21.9;26.2]	0.709	
BMI (>30 kg/m <sup>2</sup> )	24 (6.2%)	538 (5.6%)	0.692	
Age	58.0 [48.0;68.0]	60.0 [50.0;69.0]	< 0.001	
Age (>50 years)	250 (64.1%)	7324 (75.6%)	< 0.001	
Neck circumferential (>36.3 cm)	163 (41.8%)	3407 (35.2%)	0.008	
Sex (male)	186 (47.7%)	4179 (43.1%)	0.083	
Smoking	191(49.2%)	3807 (39.6%)	< 0.001	
Alcohol				
Regular user	161 (41.4%)	4358 (45.1%)		
Non-user	227 (58.4%)	5275 (54.6%)		
Comorbidity		1		
Asthma	34 (9.4%)	273 (3.0%)	< 0.001	
AR	179 (49.7%)	1000 (11.0%)	< 0.001	
House income				
High	90	2549		
Middle-high	103	2393		
Middle-low	95	2424		
Low	101	2281		
Economic activity				
Employed	196 (54.6%)	5335 (58.8%)		
Unemployed	163 (35.4%)	3742 (41.2%)		
Educated level				
High (>9 years)	263 (73.3%)	5939 (65.5%)		
Low (≤9 years)	96 (26.7%)	3135 (34.5%)		

**Table 1.** Demographics and STOP-Bang questionnaire for study subjects with and without chronicrhinosinusitis. CRS chronic rhinosinusitis, OSA obstructive sleep apnea, AR allergic rhinitis.

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the CRS group, while alcohol use was not significantly (P=0.161) different between the two groups. Asthma and AR were significantly more prevalent in the CRS group (Asthma: 9.4% of CRS patients and 3.0% of without CRS subjects; AR: 49.7% of CRS patients and 11.0% of without CRS subjects). There were no significant differences in house income (P=0.212) or economic activity (P=0.128). However, education level was higher in the CRS group (73.3% of CRS patients are highly educated level, compared to 65.5% of subjects without CRS; P=0.003).

#### Relationship between OSA risk and CRS state or symptoms

In this study, those with STOP Bang scores of 0–2, 3–4, and 5–8 were categorized into low, intermediate, and high OSA risk groups, respectively. Based on this categorization, we found that all parameters of STOP-Bang questionnaire increased significantly in a stepwise manner (Supplementary Table 1). Moreover, the presence of CRS significantly increased as the risk of OSA increased from low to high (3.1%, 3.8%, and 5.3%, respectively; P < 0.001; Table 2). There were trend differences in the timing of CRS diagnosis among the groups (median age: 39.0, 41.0, and 35.0 years; P = 0.054). However, there was no significant difference in the presence of nasal polyps among the three OSA risk groups (P = 0.463). Additionally, there was no significant difference in whether individuals were currently on treatments (P = 0.302).

In the analysis of CRS symptoms, no significant association was found between the STOP-Bang classification and anterior/posterior nasal drip or presence of facial pain (P=0.131 or P=0.803, respectively, Table 3). However, a notable relationship was observed between higher-risk OSA groups and increased likelihood of experiencing symptoms such as nasal congestion (low OSA risk group: 4.1%, intermediate risk OSA group: 5.7%, high risk OSA group: 7.3%; P<0.001) and reduced sense of smell (low risk OSA group: 1.9%, intermediate risk OSA group: 3.3%, high risk OSA group: 3.3%; P<0.001).

CRS state	Total N	Low	Intermediate	High	P value
Patients with CRS	10,081	169/5475 (3.1%)	137/3630 (3.8%)	52/976 (5.3%)	< 0.001
Diagnosis of nasal polyps	498	46/248 (18.5%)	42/179 (23.5%)	15/71 (21.1%)	0.463
Age at diagnosis of CRS	679	39.0 [23.0;49.0]	41.0 [23.0;55.0]	35.0 [17.0;52.5]	0.054
Currently receiving treatment for CRS	683	23/340 (6.8%)	27/263 (10.3%)	7/80 (8.8%)	0.302

 Table 2.
 Chronic rhinosinusitis states with high risk for OSA according to the STOP-Bang questionnaire. CRS

 Chronic rhinosinusitis. Method: Cochran-Armitage trend test, Kruskal–Wallis rank sum test.

Symptoms related with CRS	Low (n = 5475)	Intermediate (n = 3630)	High (n=976)	P value
Anterior/posterior nasal drip for more than 3 months	373 (6.8%)	275 (7.6%)	80 (8.4%)	0.131
Nasal obstructions for more than 3 months	225 (4.1%)	206 (5.7%)	71 (7.3%)	< 0.001
Facial pain or pressure for more than 3 months	25 (0.5%)	18 (0.5%)	6 (0.6%)	0.803
Anosmia or hyposmia for more than 3 months	106 (1.9%)	120 (3.3%)	32 (3.3%)	< 0.001

**Table 3.** Chronic rhinosinusitis symptoms with high risk for OSA according to the STOP-Bang questionnaire.*CRS* chronic rhinosinusitis; Method: Cochran-Armitage trend test.

#### Discussion

The goal of this study was to examine the association between CRS and OSA risk through an analysis of the relationship between the CRS and STOP-Bang questionnaire based on the KNHANES. We confirmed that subjects with CRS had higher STOP-Bang questionnaire scores through a national survey, thereby providing real-world evidence from the general population. Previous studies have reported that OSA and sleep disruption are highly prevalent in patients with CRS<sup>14,15</sup>. A study on World Trade Center Responders reported CRS is an independent risk factor for OSA, with an odds ratio (OR) of 1.80<sup>14</sup>. Furthermore, these studies have examined the relationship between CRS and OSA using internally developed sleep quality questionnaires or home sleep testing. However, to the best of our knowledge, no studies have demonstrated the relationship between CRS and the STOP-Bang questionnaire using a large nationwide database.

Additionally, our study is the first to investigate specific symptoms of CRS in relation to OSA, while also identifying which components of the STOP-Bang questionnaire are associated with CRS in the general population. Among the four symptoms of CRS (nasal congestion, discharge, facial pain/pressure, and decreased sense of smell), nasal congestion and decreased sense of smell were increased in the high risk group. According to a previous study, participants with these typical CRS symptoms had an increased risk of various sleeprelated problems such as snoring, difficulties in initiating sleep, difficulties in maintaining sleep, early morning awakening, and excessive daytime sleepiness compared to those without CRS symptoms<sup>15</sup>. Interestingly, within the STOP-Bang questionnaire, we found positive correlations of CRS with snoring, tiredness, and observed apnea. This finding suggests that airflow disturbance can increase the risk of OSA in CRS patients, providing additional information beyond previous studies that suggest a potential link between the two conditions through shared systemic inflammatory pathophysiological mechanisms<sup>9</sup>. We did not observe significant differences in hypertension, BMI, or sex among components of the STOP-Bang questionnaire for CRS patients. Similarly, a previous study has shown that patients with CRS tend to exhibit a higher risk of OSA even with lower BMI levels<sup>23</sup>. Our data showed that patients with CRS were relatively younger in age, despite the well-established association between increasing age and a higher risk of OSA<sup>24</sup>. Nevertheless, CRS patients still had a higher risk of OSA.

CRS is a heterogeneous multifactorial disease<sup>25</sup>. It is known to be associated with microbial infections, smoking, pollutants, allergens, and other social factors such as a correlation with low socioeconomic status<sup>26,27</sup>. However, in our study, it was not related to household income or economic activity. The prevalence of CRS was higher in the group with higher education. Regarding comorbidities of CRS, previous research has indicated that the presence of persistent AR in addition to CRS can further amplify the risk of sleep problems<sup>15</sup>. Moreover, conditions that are well-known risk factors for OSA, such as AR and asthma, were found to be more prevalent in patients with CRS. The presence of these conditions, which are frequently observed in CRS, is considered to increase the risk of OSA. CRS is classified into type 2 and non-type 2 according to endotypes<sup>28</sup>. In Korea, there has been an eosinophilic shift, and the prevalence of type 2 CRS is increasing<sup>29,30</sup>. Comorbidities such as AR and asthma are also on the rise, which may increase the risk of OSA.

The association between CRS and OSA is complex. Although such association is not fully established yet, some studies have suggested that these two conditions might be bidirectionally related. As mentioned earlier, CRS has been identified as a risk factor for OSA. However, some studies also suggest that OSA may increase the development of CRS<sup>31</sup>. According to Garvey et al., patients with both CRS and OSA show higher rates of antibiotic and steroid usage than those with CRS alone<sup>31</sup>. In particular, among the group of patients with both OSA and CRS who did not receive continuous positive airway pressure or sleep surgery, the likelihood of using

antibiotics or steroids remained higher even when compared to groups receiving similar rates of endoscopic sinus surgery<sup>31</sup>.

A limitation of our study was that it was designed as a cross-sectional study, making it difficult to establish causal relationships. However, in our research, CRS treatment status did not significantly correlate with the OSA risk. Additional study is needed to further investigate this. Another limitation of our study was that the KNHANES included the STOP-Bang questionnaire only for individuals aged 40 and above. Consequently, our study was limited to participants within this age group, and the findings may not be representative of all age groups. Additionally, we did not diagnose OSA through objective tests but relied on a questionnaire-based approach. To enhance the reliability of our findings, we utilized various indicators from multiple questionnaires. Furthermore, since this study relied on survey data, defining CRS through endoscopic or CT findings was not possible, nor was obtaining nasal tissue for endotyping. Future prospective studies should incorporate endoscopic examinations, CT scans, and tissue sampling from participants to address these limitations.

## Conclusions

Our study showed that the subjects with CRS had significantly higher STOP-Bang questionnaire score. This finding was associated with increased occurrences of snoring, tiredness, and apnea. Among symptoms of CRS, nasal obstruction and decreased sense of smell were found to be linked to an increased risk of OSA.

## Methods

#### **Ethics statement**

The 2019-2021 KNHANES was approved by the Institutional Review Board (IRB) of the Korea Disease Control and Prevention Agency (IRB No. 2018-01-03-C-A, 2018-01-03-2C-A, and 2018-01-03-3C-A). All participants signed informed consent forms during the survey. This study was also approved by the IRB of Soonchunhyang University Cheonan Hospital (IRB No. 2023-09-056). The study was performed in accordance with the approved guidelines and the Declaration of Helsinki.

#### Study population

This was a cross-sectional study based on nationwide population data. It was conducted using the Korean National Health and Nutrition Examination Survey (KNHANES) data from 2019 to 2021. KNHANES is an annual national survey project assessing the health and nutrition status of Korean since 1998<sup>22</sup>. It is carried out by the Korea Disease Control and Prevention Agency<sup>22</sup>. Each year, KNHANES samples approximately 10,000 nationally representative individuals using a multi-stage clustered probability design<sup>22</sup>. The KNHANES data were obtained from the official KNHANES website (https://knhanes.kdca.go.kr/knhanes). A flow chart of the participant inclusion and exclusion criteria is provided in Fig. 1. Individuals who completed both the STOP-Bang and CRS-related questionnaires were included in the study, while those who did not complete either questionnaire were excluded.

#### STOP-Bang questionnaire

The STOP-Bang questionnaire was conducted for individuals aged 40 years and older in KNHANES after the year 2019 to assess the risk of obstructive sleep apnea (OSA). The STOP-Bang questionnaire was utilized as the assessment tool. It consisted of eight items, namely: (1) Snoring: Do you snore loudly (louder than talking or loud enough to be heard through closed doors)?; (2) Tired: Do you often feel tired, fatigued, or sleepy during daytime?; (3) Observed: Has anyone observed you stop breathing during your sleep?; (4) Pressure: Do you have or are you being treated for high blood pressure?; (5) BMI > 30 kg/m<sup>2</sup>; (6) Age > 50 years old; (7) Neck circumference > 36.3 cm; and (8) Gender (if male)<sup>32</sup>. The risk of OSA was categorized based on the STOP-Bang score, with the following classifications: (1) high risk of OSA, affirmative responses to 5–8 items; (2) intermediate risk of OSA, affirmative responses to 3–4 items; and (3) low risk of OSA, affirmative responses to 0–2 items. The total count of "yes" responses was calculated as the sum of affirmative answers across all questionnaire items<sup>33</sup>.

#### Chronic rhinosinusitis and clinical factor measurements

CRS was defined based on the survey questionnaire as the presence of two or more symptoms persisting for at least 12 weeks, with at least one being nasal blockage/obstruction/congestion or nasal discharge (anterior/ posterior nasal drip)<sup>28</sup>. Optional symptoms included facial pain/pressure and reduction or loss of smell<sup>28</sup>. The survey questionnaire included items related to CRS symptoms such as anterior/posterior nasal drip, nasal obstruction, facial pain or pressure, and anosmia or hyposmia persisting for more than three months. To address potential errors, a comprehensive analysis was conducted on relevant survey items, including questions about the timing of CRS diagnosis, ongoing treatment for CRS, and the presence of nasal polyps.

Additional clinical factors included in the analysis were asthma, allergic rhinitis, smoking status, alcohol usage, household income, economic activity, and educational level. Alcohol usage was defined as regular if individuals consumed one or more drinks per month in the past year, while non-users were defined as those who had never consumed alcohol or had consumed less than one drink per month in the past year.

#### **Statistical analysis**

Categorical variables are presented as N (%). They were assessed using a chi-square test for two-group comparisons. For three or more groups, a Cochran-Armitage trend test was employed to identify any observed trends. Continuous variables are presented as median [Q1; Q3] values. If the number of data points was less than 5000, a Shapiro–Wilk Normality test was performed to assess normality. If the number of data points exceeded

5000, an Anderson-Daring test was utilized. Following the normality test, two-group comparisons utilized either a *t*-test or Mann–Whitney test based on the normality assumption, while comparison between three or more groups employed either Analysis of Variance (ANOVA) or Kruskal–Wallis Rank Sum Test depending on the normality assumption. A *P*-value of less than 0.05 was considered statistically significant. All statistical analyses and data visualization were conducted using SAS version 9.4 for Windows (SAS Institute Inc., Cary, NC, USA) and R (version 4.1.2; R Foundation for Statistical Computing, Vienna, Austria).

#### Data availability

All the data generated and/or analyzed during the current study are included in this article and are available from the Korea National Health and Nutrition Examination Survey homepage (https://knhanes.kdca.go.kr/knhanes/sub03/sub03\_02\_05.do).

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# Author contributions

Conceptualization and study design: HKC, JHC; data acquisition: HKC, DK, HWL, YL; Statistical analysis and visualization: HKC; the original draft of the manuscript: HKC, JHC; supervision or mentorship: BB, JYL, JHC. All authors reviewed the manuscript.

# **Competing interests**

The authors declare no competing interests.

# Additional information

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