

Assessment of Change in Palatal Sensation in Obstructive Sleep Apnea Patients by Using Two-Point Palatal Discrimination

Kyung-Hwa Jeong¹ · Youngsoo Yang¹ · Hye Rang Choi¹ · Jae Hoon Cho¹ · Gee-Tae Kim² · Jin Kook Kim¹

Departments of ¹Otorhinolaryngology-Head and Neck Surgery, ²Biomedical Engineering, Konkuk University School of Medicine, Seoul, Korea

Objectives. Patients with obstructive sleep apnea syndrome (OSAS) have impaired responses to inspiratory resistive loading during sleep. This may be due, in part, to a change in the upper airway sensation. Therefore, we hypothesized that patients with OSAS have diminished upper airway sensation due to snoring.

Methods. A total of 53 participants were selected based on clinical evaluation and polysomnography. Two-point discrimination was measured with modified calipers in the tongue and soft palate.

Results. A total of 10 participants were included in the control group, 12 participants in the simple snoring group, and 27 participants in the OSAS group. There were 12 patients in the impaired sensation group of the OSAS group. On comparing polysomnography, patients with impairment of their palatal sensory input in two-point discrimination (TPD) had a more protracted duration of the longest snoring episode than those with simple snoring and normal sensation. Patients with decreased sensory input in TPD had longer average duration of snoring episodes and relative snoring time than those with simple snoring and normal sensory input in cold uvular TPD. Comparison of the cold uvular TPD for normal sensation and impaired sensation in patients with OSAS after treatment showed a different trend.

Conclusion. Impaired sensation of the soft palate was correlated with the longest snoring episode duration, average snoring episode duration, and relative snoring time. It is helpful in detecting the early stage of neural degradation in OSAS patients by assessing snoring components of polysomnography and TPD in the soft palate.

Keywords. Sensory Neuropathy; Sleep Apnea Syndromes; Snoring; Polysomnography; Palate, Soft

INTRODUCTION

Apneic episode in obstructive sleep apnea (OSA) starts with the collapse of the upper airway during sleep, and resolves when stimuli generated during apneas cause a brief arousal and re-opening of the airway. In general, obstructive sleep apnea syndrome (OSAS) is progressive over time and its prevalence increases with age [1,2]. Svanborg [3] proposed and pursued the

hypothesis on the pathogenesis of OSA that long-standing snoring-induced vibrations cause neurogenic lesions in upper airway tissues, and then cause damage to the reflex circuits which keep the upper airway open during inspiration [3]. This could be critical during sleep because the muscle tone is normally reduced. Several studies have supported this hypothesis by confirming upper airway neuropathy in OSAS patients [3-5]. Other research groups have also supported the hypothesis with contributing data [6-8]. In these studies, various methods for measuring local sensory neuropathy have been used, such as vibration [6], 2-point discrimination [7], and air-pressure pulses [8]. However, no attempts have been made to correlate the degree of sensory deficit in the soft palate with the duration of snoring.

We have demonstrated measurement of warm and cold two-point detection (TPD) with the noninvasive method of levels (MLE). In the MLE, stimuli of predetermined intensities are

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• Corresponding author: **Jin Kook Kim**

Department of Otorhinolaryngology-Head and Neck Surgery, Konkuk

University Medical Center, Konkuk University School of Medicine,

120-1 Neungdong-ro, Gwangjin-gu, Seoul 05030, Korea

Tel: +82-2-2030-7662, Fax: +82-2-2030-5299

E-mail: entalk@kuh.ac.kr

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used and the test subject responds after the stimulus has been given to confirm whether it was detected or not. The purpose of this study was to use this method to compare sensory deficit in the soft palate among untreated subjects with OSAS, and to correlate these data with the apnea-hypopnea index (AHI) and the duration of snoring.

MATERIALS AND METHODS

Subjects

Patients with sleep disorders and normal subjects who fulfilled the following clinical and polysomnographic inclusion criteria were included: (1) clinical criteria, age between 17 years and 65 years (or premenopausal women); absence of neurologic, cardiovascular, pulmonary, or other chronic illness; no prior surgery of the nose and palate; no current drug intake; modified Friedman stage I; oropharyngeal obstruction during the Müller maneuver; and no prior treatment of sleep-disordered breathing; (2) polysomnographic recording criteria, patients with OSAS should have an AHI ≥ 5 /hr of sleep; however, simple snorers should have an AHI < 5 /hr of sleep with concomitant snoring (relative snoring time $\geq 5\%$) on polysomnography; otherwise, the control group should have an AHI < 5 /hr of sleep without a snoring event (relative snoring time $< 5\%$) during polysomnography; and (3) subjects should provide informed consent. Exclusion criteria were any previous treatment for snoring or OSA, previous upper airway surgery excluding remote tonsillectomy, previous cerebrovascular accident, diabetes, any neuropathy or active neurologic disease, and recent upper respiratory tract infection, excessive gag reflex, or a very small oropharyngeal cavity at the time of proposed testing, which precluded sensory testing. All OSA patients underwent lateral pharyngoplasty and tonsillectomy with sparing of the uvula. We evaluated the initial TPD on the same day the patient underwent polysomnography before surgery. Also, the TPD during follow-up was measured after 3 months of surgery. Approval for the trial was granted by Konkuk University Hospital Ethics Committee (KUH1110017).

Polysomnography

All subjects with an initial evaluation who met the inclusion cri-

teria underwent nocturnal polysomnography. The following variables were monitored: electroencephalogram (EEG), electro-oculogram, chin electromyogram, leg electromyogram, heart rate (modified V₂ lead), and body position. Respiration was evaluated with nasal cannula pressure transducer system, polygraphic recording with esophageal pressure (PES) transducer, thoracic bands, abdominal bands, pulse oximeter, and neck microphone. Recordings were scored following the international criteria for sleep/wake with identification of short (≥ 3 seconds) arousals; tabulation of apneas, hypopneas; and obstructive, mixed events, and central events following the criteria of the American Academy of Sleep Medicine [9]. There was also scoring of short visual arousals (≥ 3 seconds) secondary to 'abnormal breathing efforts.' This scoring was performed using PES recording with recognition of 'PES crescendos' and 'sustained abnormal respiratory effort' as previously described [10], in addition to the nasal cannula/pressure transducer system with recognition of respiratory events of < 10 seconds duration but terminated by a visual EEG arousal of ≥ 3 seconds (arousal with abnormal breathing effort). Snoring components consisted of relative snoring time (%), average snoring episode duration (minutes), longest snoring episode (minutes), total snoring time (minutes), and number of snoring episodes. These components were calculated on polysomnography (Somnologica Studio, Embla Systems Inc., Broomfield, CO, USA).

Palatal test

A special compass-like device was built by one of the authors (GTK) (Fig. 1). When the two branches were in close contact, the distance was 0.2 mm in diameter. A screw system allowed opening of the two branches to keep the two branches fixed at the selected distance. A thermode was applied to the soft palate and the anterior tongue. TPD was determined with the MLE [11]. The apparatus was set at a starting temperature of 43°C and 0°C for soft palate testing and anterior tongue testing, respectively.



Fig. 1. Modified caliper device. Special compass-like device consisting of temperature display and control buttons (A), scale (B), and thermode with a screw system (arrow).

HIGHLIGHTS

- Patients with impaired palatal sensory input in two-point detection had more protected duration of the longest snoring episode.
- Those also had longer average duration of snoring episodes and relative snoring time.
- Impaired palatal sensation was well correlated with all the snoring measurements.

Then, each subject was asked to indicate 1 or 2 points by holding up the appropriate number of fingers to confirm whether or not he/she perceived a sensation of cold (warm) as soon as they perceived a sensation of cold (warm) [12]. The first stimulus step size was 6 mm. If the subject did not perceive the first stimulus, the next stimulus was increased in the 3-mm size until the subject responded (first stimulus 6 mm, second stimulus 9 mm, third stimulus 12 mm etc.). If the subject first perceived a stimuli sensation at for example 6 mm, new stimuli, with subtraction of 2-mm step size from the previous round, were repeated until a 'no' response was obtained (6, 4, 2, 1 mm etc.). If the subject then perceived stimuli at for example 1.0 mm, the intensity of the next stimulus was increased again in steps half the size of those in the previous round (1, 2, 3 mm etc.) until a 'yes' response was obtained. The time intervals between the four stimuli were randomized to last between 4 and 6 seconds. The length of each test was measured by one of the authors.

As there was no available definition of 'impaired sensation of the soft palate,' we defined it as the length of discrimination distance of the soft palate is longer than one-half of the length of discrimination distance of the tongue individually.

Statistical analyses

Descriptive statistics were used for population and group characteristics. Correlations between soft palate TPD and age, AHI, and snoring components of polysomnography were evaluated with group analysis performed using Kruskal-Wallis test. Receiver operating characteristic (ROC) curve was applied for the cutoff value of sensory change [13]. To compare between the pre-treatment TPD and post-treatment TPD, the Wilcoxon signed rank test was performed. All data were analyzed with IBM SPSS statistical software ver. 20.0 (IBM Co., Armonk, NY, USA). Statistical significance was set at $P < 0.05$.

RESULTS

Demographics of participants

A total of 53 individuals were included in the study. Two subjects (two men) could not participate in TPD testing due to strong gag reflexes, and two subjects (both men) had technically unsatisfactory recordings. Characteristics of the remaining forty-two men and seven women included in the analysis are summarized in Table 1. The age and body mass index (BMI) were significantly different in the control group, but no difference was observed between the simple snoring group and the OSAS group, and AHI in the OSAS group was higher than that in the other groups. Compared with the control group, the mean TPD distance at 0°C and 43°C was longer in the OSAS group, but it was not significantly different. There were no significant differences in age and TPD between the OSAS group and the control group.

Formation of the impaired sensation group

Some subjects in the OSAS group displayed longer TPD distance than the other subjects in the same group. Furthermore, their snoring components tended to increase. We investigated the cutoff value for defining the sensory change. The mean length of 2.5 mm in the soft palate cold TPD test obtained by the ROC curve analysis was the cutoff value with 91.7% sensitivity and 85.7% specificity (area under curve=0.901, $P < 0.001$) (Table 2).

There were 15 patients in the normal sensation group and 12

Table 2. Sensitivity and specificity of cutoff point for diagnosing impaired sensation group at different thresholds

Cutoff point (mm)	Sensitivity (%)	Specificity (%)	$(1 - \text{sensitivity})^2 + (1 - \text{specificity})^2$
1.5	100	33.3	0.4444
2.5	91.7	85.7	0.0274
3.5	83.3	85.7	0.0482
5.0	33.3	99.5	0.4467

Table 1. Subject characteristics

Characteristic	Control	Simple snorer	Obstructive sleep apnea syndrome	P-value*
No. of subjects	10	12	27	
Sex (male:female)	8:2	9:3	25:2	
Age (yr)	35.3±13.2	32.8±9.7	44.6±12.7	0.029
Body mass index (kg/m ²)	22.2±2.5	22.2±1.3	27.0±2.1	<0.001
Apnea-hypopnea index	2.1±1.7	2.8±2.7	35.3±18.4	<0.001
TPD test (43°C)				
Anterior tongue (mm)	1.5±0.7	1.7±0.6	1.9±0.8	0.462
Soft palate (mm)	1.3±0.5	1.3±0.6	3.1±1.9	0.084
TPD test (0°C)				
Anterior tongue (mm)	1.5±0.5	1.7±0.6	2.2±1.0	0.697
Soft palate (mm)	1.3±1.4	1.3±0.6	2.2±1.0	0.130

Values are presented as mean±SD.

TPD, two-point discrimination.

*Kruskal-Wallis test.

patients in the impaired sensation group (Fig. 2). There were no significant differences in age, BMI, and AHI between the two groups (Table 3). The results of the 43°C TPD test in the anterior tongue and soft palate showed that the simple snoring group had a significantly shorter distance than the other groups ($P=0.004$, $P<0.001$, respectively). Also, the results of the soft palate cold TPD test demonstrated that the impaired sensation group had a significantly longer distance than the other groups ($P<0.001$).

Comparison of polysomnography

On polysomnography, the mean AHI in the OSAS group was

higher, and varying results were observed for snoring components. Average snoring episode duration, longest snoring episode, and relative snoring time in the impaired sensation group were significantly longer than those in the normal sensation group ($P=0.043$, $P=0.010$, and $P=0.032$, respectively) (Figs. 3–5) and lowest O₂ saturation was significantly different in the simple snoring group ($P=0.009$).

Alteration of cold TPD between the normal and impaired sensation groups of the OSAS group

Compared to the mean value before treatment, the mean value after treatment did not significantly change for cold TPD in the

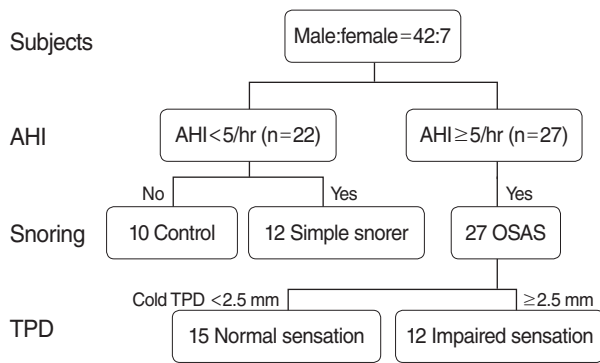


Fig. 2. Demographics of subjects. A total of 49 patients were enrolled. The number of subjects in the control group was 10, that in the simple snoring group was 12, that in the normal sensation group of the OSAS group was 15, and that in the impaired sensation group of the OSAS group was 12. AHI, apnea-hypopnea index; TPD, two-point detection; OSAS, obstructive sleep apnea syndrome.

Table 3. Snoring components between the normal sensation group and the impaired sensation group

Variable	Normal sensation group in OSAS	Impaired sensation group in OSAS	P-value*
Sex (male:female)	15:0	10:2	
Age (yr)	42.7 ± 13.4	47.0 ± 12.0	0.032
Body mass index (kg/m ²)	26.9 ± 2.2	27.2 ± 2.1	<0.001
Apnea-hypopnea index	37.9 ± 21.5	32.0 ± 13.8	<0.001
TPD test (43°C)			
Anterior tongue (mm)	1.9 ± 0.7	1.8 ± 0.8	0.004
Soft palate (mm)	3.1 ± 2.0	3.0 ± 1.8	<0.001
TPD test (0°C)			
Anterior tongue (mm)	1.9 ± 0.7	2.8 ± 1.2	0.114
Soft palate (mm)	2.1 ± 1.3	4.6 ± 1.6	<0.001

Values are presented as mean ± SD. OSAS, obstructive sleep apnea syndrome; TPD, two-point discrimination. *Kruskal-Wallis test.

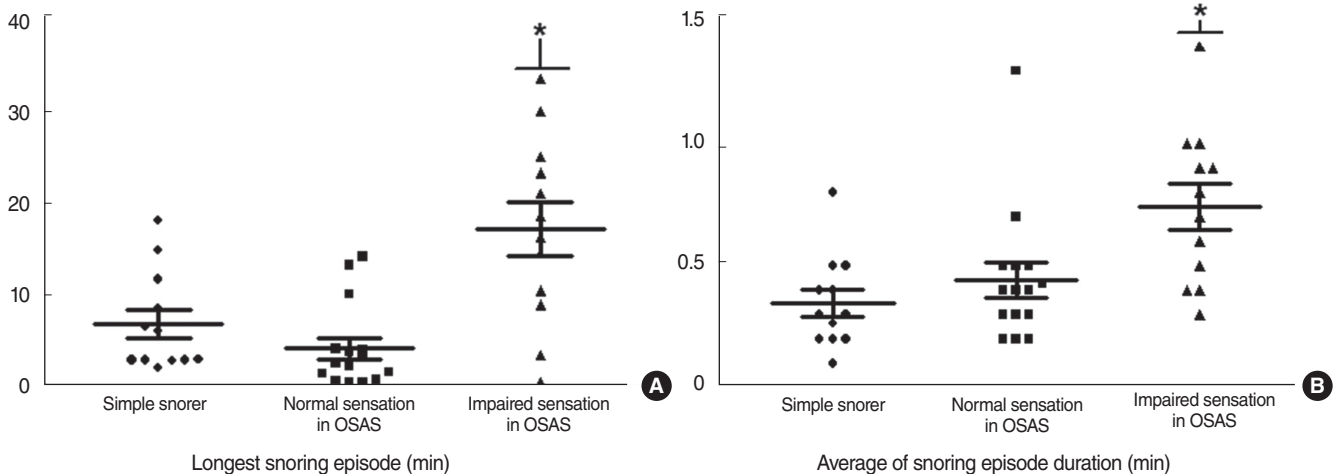


Fig. 3. (A) Longest snoring episode (min). The impaired sensation group of the OSAS group showed a significant difference. The mean ± SD was 15.6 ± 10.3 in the impaired sensation group, 7.4 ± 5.6 in the simple snoring group, and 3.9 ± 4.5 in the normal sensation group of the OSAS group. Symbol (*) indicates statistical significance. (B) Average snoring episode duration (minute). The impaired sensation group of the OSAS group showed a significantly longer duration than the other groups. The mean ± SD was 0.7 ± 0.4 in the impaired sensation group, 0.4 ± 0.2 in the simple snoring group, and 0.4 ± 0.3 in the normal sensation group of the OSAS group. Symbol (*) indicates statistical significance. OSAS, obstructive sleep apnea syndrome.

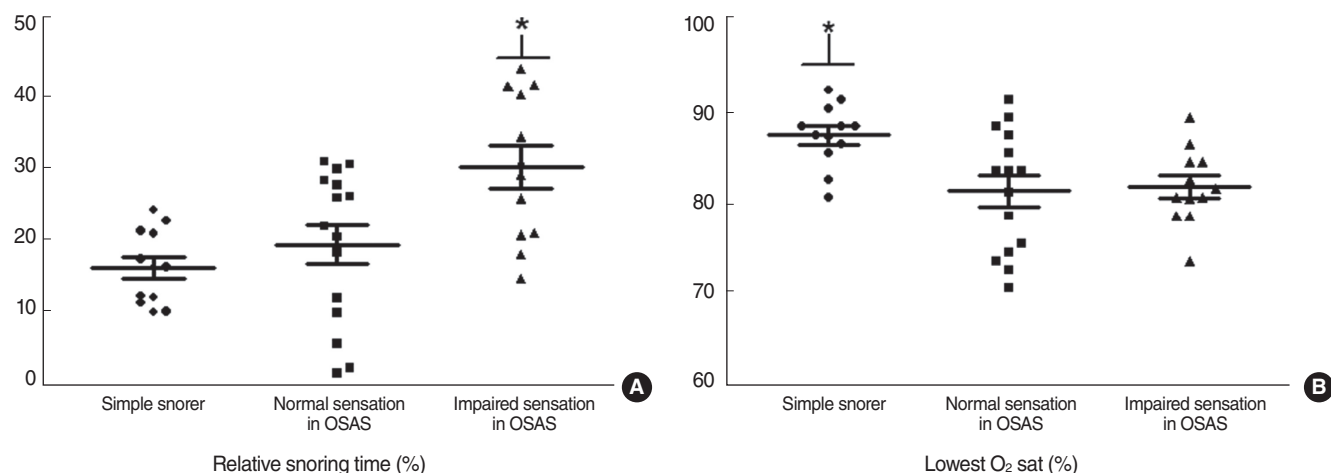


Fig. 4. (A) Relative snoring time (%). The mean \pm SD was 16.8 ± 4.8 in the simple snoring group, 18.2 ± 11.3 in the normal sensation group, and 28.7 ± 10.8 in the impaired sensation group of the OSAS group. The impaired sensation group showed a significant difference. Symbol (*) indicates statistical significance. (B) Lowest O₂ saturation (%). The mean \pm SD was 86.9 ± 3.5 in the simple snoring group, 80.7 ± 6.8 in the normal sensation group, and 79.9 ± 6.5 in the impaired sensation group of the OSAS group. There was no significant difference between the normal and impaired sensation groups except in the simple snoring group. Symbol (*) indicates statistical significance. OSAS, obstructive sleep apnea syndrome.

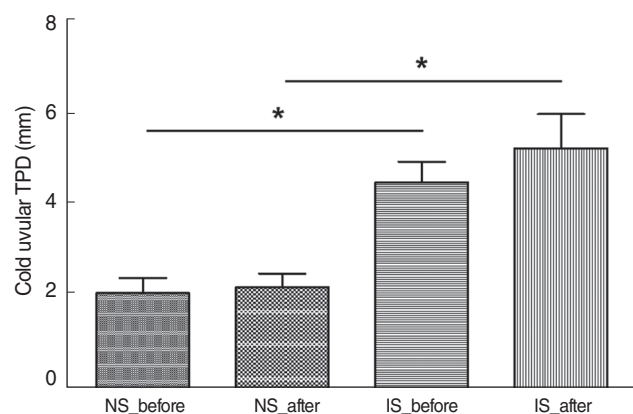


Fig. 5. Alteration of cold TPD between the normal and impaired sensation groups of the OSAS group. Compared to the mean before treatment, the mean after treatment in the normal sensation group did not change significantly for cold TPD ($P=0.291$). In the impaired sensation group, the cold uvular TPD before treatment was increased without statistical significance ($P=0.392$). Symbol (*) indicates statistical significance. NS_before, the normal sensation group before treatment; NS_after, the normal sensation group after treatment; IS_before, the impaired sensation group before treatment; IS_after, the impaired sensation group after treatment; TPD, two-point detection; OSAS, obstructive sleep apnea syndrome.

normal sensation group ($P=0.291$) (Fig. 5). In the normal sensation group, the mean \pm SD before treatment was 2.1 ± 1.3 , and the mean \pm SD after treatment was 2.3 ± 1.2 . In the impaired sensation group, the mean \pm SD before treatment was 4.6 ± 1.6 , and the mean \pm SD after treatment changed to 5.3 ± 2.6 without statistical significance ($P=0.392$). But, the difference between the normal and impaired sensation groups in cold TPD after treatment was statistically significant ($P=0.001$).

DISCUSSION

Segmental demyelination and axonal degeneration of afferent neurons, such as α -fibers, lead to sensory impairment. These lesions are responsible for slowing of impulse conduction [14]. Friberg et al. [15,16] provided evidence for local neurogenic lesions in heavy snorers and OSAS patients [3]. Other studies supported the hypothesis that snoring is associated with the histologic alterations; Kimoff et al. [6] suggested abnormal responses in snorers. Our data partially agree with these studies. The patients in the simple snoring group did not show any sensory change in the soft palate compared to those in the control group, but the patients in the OSAS group showed sensory change in the soft palate.

On comparing TPD in cold and warm temperatures, there was no significant difference between the OSAS group and the control group. This finding is similar to that in the report by Guilleminault et al. [7], although we did not separate the upper airway resistance syndrome subjects and the OSAS subjects. The previous authors explained that there was not enough snoring during the night and/or the number of years spent snoring was insufficient in patients with OSAS to induce the neurologic lesions. In contrast to the present study, Sunnergren et al. [17] reported a significant positive correlation between both estimated periods of snoring and they objectively evaluated the degree of sleep-disordered breathing with the degree of impaired sensation of the soft palate at an individual level. However, in this prior study, snoring was not recorded objectively.

Our study strengthens the hypothesis that long-standing, snoring-induced vibrations could cause impaired sensation of the soft palate, and that sensory change in the soft palate might

be involved in the pathologic progression often seen in OSA. It is well-known that long-term exposure to low-frequency vibration causes peripheral nerve injury in humans [18]. In our study, we determined 2.5 mm in cold TPD as the cutoff value for impaired sensation of the soft palate, which could possibly be used for early detection of palatal peripheral neuropathy in OSAS subjects. Previous studies only made a comparison among non-snorers, snorers, and OSA subjects [6,7,17] without the use of a cutoff value. In our study, there was no difference between pre-operative TPD and postoperative TPD. Therefore, further studies with long-term observation and larger sample size are needed to verify our suggestion because a short follow-up period and a small sample size were used in this study.

Long-standing vibrations not only cause sensory neuropathy due to thermal stimuli but also due to mechanical stimuli [19,20]. In agreement with this phenomenon, an interesting finding of the present study was that the OSAS patients with greater longest snoring time, average snoring episode duration, and relative snoring time on polysomnography tended to have impaired sensation of the soft palate. This finding was also consistent with previous studies [6,7,15-17]. In spite of the conception that the subocclusive stage in snoring precedes the development of OSA [21], there was a broad spectrum ranging from mild to severe snorers among OSAS subjects. This indicated that it was not always a one-way progression from snoring to OSA. We speculate that vibration-induced snoring may have played a more important role in sensory change in the soft palate than oxygen desaturation in OSAS subjects. The clinical importance of sensory change in the soft palate was not totally deciphered in the present study, but based on the concept that the continuous positive airway pressure (CPAP) treatment created airflow buttress that reduced snoring, CPAP would be helpful in preventing impaired sensation of the soft palate.

The limitations of this study were its small sample size and the lack of comparison of clinical symptoms. Long-term, follow-up observations are needed. The limitation of the present study also included unknown duration of OSAS because subjects could not report the accurate time of onset of symptoms. A further study assessing the change in snoring components after treatment is needed. Another limitation that should be considered is that psychophysical methods, such as two-point discrimination testing, require active participation of the subjects. Therefore, the results could have been affected by psychological confounding factors. Also, a comparison of the tools used in this study and the previous study was not performed. Although the devices used in the previous study had a relatively same scale range as our device, a further study comparing the published tools is needed to determine the accuracy of TPD.

In conclusion, impaired sensation of the soft palate is correlated with longest snoring episode duration, average snoring episode duration, and relative snoring time, and hence, it could be useful in detecting the early stage of neural degradation in OSAS

patients through assessment of snoring components of polysomnography and two-point discrimination in the soft palate.

CONFLICT OF INTEREST

No potential conflict of interest relevant to this article was reported.

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