#### ORIGINAL RESEARCH

## High Tongue Position is a Risk Factor for Upper Airway Concentric Collapse in Obstructive Sleep Apnea: Observation Through Sleep Endoscopy

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**Purpose:** Identification of upper airway (UA) obstruction based on pharyngeal factors is important for obstructive sleep apnea (OSA) evaluation. This study is to assess the association between UA collapse characteristics and Friedman tongue position (FTP) in patients with OSA through drug-induced sleep endoscopy (DISE).

**Patients and Methods:** Retrospective study in individuals with OSA who were intolerant to continuous positive airway pressure (CPAP) treatment, submitted to DISE between June 1, 2013, and July 31, 2017. All subjects were classified as having an FTP grade of I to IV, and the velum, oropharynx, tongue base, epiglottis (VOTE) classification was used to analyze the DISE findings. UA collapse characteristics by DISE and FTP grading were compared between groups. The associations between specific DISE findings and FTP were analyzed.

**Results:** In total, 205 patients were assessed. A positive and significant correlation was identified between the presence of retropalatal complete concentric collapse (CCC) and FTP grade, according to the following distributions: I, 17.4%; II, 22.9%; III, 33.7%; and IV, 48.7% (P = 0.014). A logistic regression model revealed that CCC was associated with FTP grade IV. After adjusting for age, sex, body mass index (BMI), and tonsil size (TS), the grade IV individuals had a 4.4-fold higher risk of having CCC than grade I individuals (P = 0.026). Multiple collapse sites and palatopharyngeal or combined (palatopharyngeal and hypopharyngeal) collapse were more prevalent in grade IV individuals.

**Conclusion:** OSA patients intolerant to CPAP have a strong positive correlation between the FTP grade and presence of retropalatal CCC. FTP grade IV is an independent risk factor for velum-CCC, controlling for sex, age, BMI, and TS grade.

**Keywords:** drug-induced sedation endoscopy, sleep endoscopy, obstructive sleep apnea, tongue position, upper airway collapse, complete concentric collapse

#### Introduction

Obstructive sleep apnea (OSA) is a common sleep disorder with a prevalence of 2% to 37% and 4% to 50%, respectively, in males and females, based on a review of 11 epidemiological studies published between 1993 and 2013, effectively showing that its prevalence has been rising in recent years.<sup>1</sup> OSA patients present typical symptoms and morbidities such as daytime sleepiness, nocturnal snoring, wheezing, increased breathing effort, frequent awakenings, and intermittent hypoxemia that compromise their sleep quality, alongside cognitive and behavioral impairment, as well as increased mortality.<sup>2,3</sup>

Upper airway (UA) obstruction during sleep is the main pathogenic mechanism underlying OSA. Several factors contribute to airway collapse susceptibility, such

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as age, gender, body mass index (BMI), neck and waist circumference, craniofacial development and pharyngeal cavity anatomy.<sup>4</sup> Among the structures that may contribute to this obstruction, the tongue is one of the most important for pharyngeal collapse, due to its increased volume, a compromising tonus, and its interaction with surrounding tissues. By cephalometric analysis, the tongue was found to contribute to velopharyngeal obstruction, as a higher tongue base resulted in increased contact distance with the soft palate; and in hypopharyngeal obstruction, as a more inferior hyoid resulted in an upright tongue position with more tongue tissue occupying the airway.<sup>5</sup> Based on the relative tongue position, it is possible to predict risk and severity of OSA and sites of UA obstruction.<sup>6–8</sup> A higher tongue position has already been associated with a higher apnea-hypopnea index (AHI), markedly severe hypopharyngeal obstruction,<sup>9</sup> and a lower surgical success rate of uvulopalatopharyngoplasty (UPPP).<sup>10,11</sup>

Two classification systems have already addressed the assessment of tongue height based on physical examination of the oral cavity, but not on tongue size. The first system is the modified Mallampati classification (MMC). Initially described as the Mallampati classification,<sup>12</sup> its score varies from class I to III based on the visualization of the uvula, tonsils, pillars and soft palate with tongue projection. In 1987, Samsoon and Young proposed a modification of the scoring system by adding a class IV, allowing for more details in describing the palate position in relation to the tongue.<sup>13</sup> This classification has been used as a clinical predictor of successful intubation in anesthesiology.<sup>8</sup> The second classification system is the Friedman tongue position (FTP) grade, reported by Friedman in 1999. FTP allows for the description of the tongue's natural position on the mouth floor and yields a score varying from I to IV, similar to the MMC.<sup>14</sup> Both have proven useful in assessing those with OSA.<sup>15</sup> However, the FTP better reflects OSA severity than the MMC,<sup>16</sup> and it is considered a classic indicator of OSA, while effectively informing surgical selection.<sup>17</sup>

Although many studies have identified a strong association between an increased OSA incidence and severity and an increased score in the MMP/FTP classifications,<sup>9–11,18</sup> the clinical features correlated with FTP grades remain diverse. Harvey et al showed that individuals with high FTP scores presented with overall UA volume reduction, and not only a narrowing of the retroglossal area by computed tomography.<sup>19</sup> A relatively high tongue position was found to be associated with increases in soft palate length and UA obstruction.<sup>20</sup> In addition, some studies revealed no relationship between increasing FTP and UA obstruction.<sup>21,22</sup>

Drug-induced sleep endoscopy (DISE) is a tool widely used to evaluate UA collapse in OSA individuals in recent years, despite some well-known limitations.<sup>23</sup> It enables a dynamic and consecutive observation of the airway lumen during drug-induced sleep, mimicking natural sleep. Different collapse patterns have enabled accurate surgery selection and management.<sup>24</sup> These DISE findings may help explain the contradictions in the literature regarding FTP grade and OSA severity.

In this context, this study aimed to identify the characteristics of UA collapse in relation to FTP grade, especially in regard to high grades, in OSA individuals.

## Patients and Methods Study Design

This was a retrospective study that analyzed all DISE exams performed on OSA patients at the Stanford Sleep Surgery Clinic between June 1, 2013, and July 31, 2017, and was approved by the Institutional Review Board and Hospital Research Ethics Committee of Stanford Hospital and Clinics (Protocol 35054).

Individuals with an apnea-hypopnea index (AHI)  $\geq 5$  events/hour, recorded by overnight polysomnography (PSG) in the laboratory or through a home sleep test (HST), and also with continuous positive airway pressure (CPAP) intolerance, were included in the study. Individuals with a previous invasive treatment for OSA, such as UPPP, tongue base surgery (transoral robotic surgery, radiofrequency, or coblation), genioglossus advancement, or maxillomandibular advancement, were excluded.

The sample characteristics, including age, sex, body mass index (BMI), tonsil size (TS) grading, FTP grading, Epworth Sleepiness Scale (ESS) score, sleep study data such as AHI and oxygen desaturation index (ODI), lowest oxygen saturation (LSAT), and DISE findings were extracted from the Stanford Hospitals and Clinics electronic health record system (REDCap [Research Electronic Data Capture] and Epic) databases. The above-mentioned clinical features were compared between groups. The associations of UA collapse characteristics (sites, patterns and degree) and FTP grades I to IV were analyzed.

## **Sleep Studies**

All patients underwent overnight PSG or HST according to the American Academy of Sleep Medicine 2012 edition.<sup>25</sup>

Apnea was identified when the airflow amplitude decreased by  $\geq 90\%$  for  $\geq 10$  seconds. Hypopnea was identified when airflow amplitude decreased by  $\geq 30\%$  for  $\geq 10$  seconds and was associated with an oxygen desaturation increase by  $\geq 3\%$  or arousal. AHI represented the number of apnea and hypopnea events per hour. ODI represented the number of times per hour that oxygen desaturation increased by  $\geq 3\%$ . LSAT represented the lowest O<sub>2</sub> saturation during sleep.

# Friedman Tongue Position and Tonsil Size Grading

FTP grading was performed with an open mouth and the tongue resting on the mouth floor and evaluated with a minimum of 5 times. Individuals included in the study were subdivided into the following four groups: FTP I, uvula and tonsils completely visible; FTP II, visible uvula but not tonsils; FTP III, visible soft palate, but not uvula; FTP IV, visible hard palate only. Tonsil size was graded accordingly: 0, tonsils absent; 1, tonsils hidden inside the pillars; 2, tonsils extending to the pillars; 3, tonsils extending to the midline.<sup>10</sup> All evaluations were performed and graded by the same senior and experienced surgeon (R.C).

#### Drug-Induced Sleep Endoscopy

DISE was performed in the operating room in a supine position. Dexmedetomidine was used for sedation in two stages, with an initial intravenous bolus of 1.5 mcg/kg over 10 minutes, followed by a continuous infusion at 1.5 mcg/kg/h. After two consecutive cycles of snoring and stopping breathing, a flexible endoscope was inserted through the nostril and positioned in the retropalatal and retrolingual regions. The minimum time for the exam recordings was 20 minutes, reducing the chances of collapse by over-induction. The Velum-Oropharynx-Tongue base-Epiglottis (VOTE) classification was used to describe the DISE findings, according to the following characteristics: sites (velum [V], oropharynx [O], tongue base [T], and epiglottis [E]); pattern (anteroposterior [AP], lateral [L], or concentric [C]); and degree of collapse (absent [0], partial [1], or complete [2]).<sup>26</sup> The VOTE obstruction sites were also analyzed as palatopharynx (higher), velum, and/ or oropharynx; hypopharynx (lower), tongue base, and/or epiglottis; and combined, in both cases. All DISE exams were performed and classified by the same senior and experienced surgeon (R.C).

#### Statistical Analysis

Normally distributed continuous variables were represented as mean and standard deviation (SD) and compared between FTP classification groups using ANOVA. Non-normally distributed variables were represented as median (range) and compared using a Wilcoxon Rank Sum test. Categorical variables were presented as frequencies (percentages, %) and compared using a Fisher's exact test or Wilcoxon Rank Sum test if categories were ordered. Logistical regression model was used with FTP grades as the main predictor and binary collapse characteristics, including degree (no, partial, complete) and pattern (AP, L, C) at sites V, O, T and E as outcome variables. The odds ratio (OR) of collapse associated with FTP grades II, III, and IV in reference to FTP grade I were then estimated without and with adjustment for age, gender, BMI, and tonsil size. Differences were considered statistically significant if P < 0.05. Statistical analyses were performed using SAS 9.4 (SAS Inc., Cary, NC, USA).

#### Results

Among 225 records found, 20 were excluded, most commonly because these individuals had undergone a previous UA surgery. The samples (205) were allocated to FTP I (n = 23), FTP II (n = 48), FTP III (n = 95), and FTP IV (n = 39) groups based on the FTP classification, and no significant differences were found in the analyzed baseline characteristics (except for age) including gender, BMI, ESS, AHI, ODI, LSAT, and tonsil size (Table 1).

The presence of complete collapse in more than one site was the most common finding, highlighting the involvement of combined palatopharyngeal and hypopharyngeal collapse and isolated palatopharyngeal collapse. Isolated collapse of the hypopharyngeal sites (T and E) was the least commonly seen among the groups. No significant differences were identified between collapse sites and different FTP grades (Table 2).

The collapse patterns and degrees at the various sites, across different FTP grades, are shown in Table 3. The presence of velum complete concentric collapse (CCC) increased significantly among groups as the FTP grade increased: 17.4%, 22.9%, 33.7%, and 48.7%, respectively, for FTP grades I to IV (P = 0.014). Collapse differences at the tongue base and other sites between groups were not significantly different (P > 0.05).

Characteristics	FTP I (n=23)	FTP II (n=48)	FTP III (n=95)	FTP IV (n=39)	P-value
Age (y) <sup>a</sup>	43.57±13.85	51.52±15.87	50.83±14.08	42.28±16.28	0.04
Male <sup>b</sup>	20 (87%)	34 (70.8%)	79 (83.2%)	30 (76.9%)	0.268
BMI (kg/m <sup>2</sup> ) <sup>a</sup>	27.17±4.57	26.74±4.21	28.46±4.21	28.84±5.55	0.082
ESS <sup>a</sup>	8.62±5.02	9.09±5.89	10.2±5.07	9.89±4.67	
c	8 (5–13)	8 (4–14)	10 (6-14)	10 (7–13)	0.585
AHI (events/h) <sup>a</sup>	30.87±19.4	34.54±21.29	38.04±20.13	38.41±22.13	
c	28 (17.4–42.1)	30.9 (19.1-45.5)	34.5 (21.9-51.3)	39.5 (19.8–53.6)	0.396
ODI (events/h) <sup>a</sup>	22.89±23.36	20.87±16.39	27.09±23.37	25.18±22.43	
c	15.4 (3.15-32.65)	15.15 (10.5–30)	20.5 (10-41.3)	19.75 (8.85–31.15)	0.647
LSAT (%) <sup>a</sup>	84.02±9.86	84.44±5.92	82.28±8.89	84±7.9	
c	87 (82–90)	85.5 (81–89)	84 (79–89)	87 (78–90)	0.419
TS <sup>b</sup>					
Grade 0	3 (13%)	8 (16.7%)	19 (20%)	8 (20.5%)	
Grade I	14 (60.9%)	28 (58.3%)	47 (49.5%)	15 (38.5%)	
Grade 2	4 (17.4%)	10 (20.8%)	25 (26.3%)	9 (23.1%)	
Grade 3	2 (8.7%)	2 (4.2%)	4 (4.2%)	5 (12.8%)	
Grade 4	0 (0%)	0 (0%)	0 (0%)	2 (5.1%)	0.19

Table I Sample Baseline Characteristics According to Different FTP Grades

Notes: <sup>a</sup>Mean ± Standard Deviation; <sup>b</sup>n (%); <sup>c</sup>Median (Range).

Abbreviations: FTP, Friedman tongue position; I, uvula and tonsils completely visible; II, visible uvula and not tonsils; III, visible soft palate, but not uvula; IV, visible only hard palate; BMI, body mass index; ESS, Epworth Sleepiness Scale; AHI, Apnea Hypopnea Index; ODI, Oxygen Desaturation Index; LSAT, lowest oxygen saturation; TS, tonsil size; grade 0, absent; grade 1, behind tonsillar pillars; grade 2, extend to pillars; grade 3, visible beyond pillars; grade 4, enlarged to midline.

FTP grade IV was found to be significantly associated with an increased velum-CCC by a logistic regression model, compared to the FTP I group, persisting after adjustment for age, gender, BMI, and tonsil size (Table 4). The FTP IV group was about 4.5-fold more likely to have a velum-CCC than the FTP I group (P = 0.026) by an OR analysis. Other collapse characteristics were not associated with the FTP grades (P > 0.05).

### Discussion

In this study, we identified greater proportions of retropalatal CCC in OSA patients with a higher FTP grade. FTP grade IV was an independent risk factor for velum-CCC, controlling for age, gender, BMI, and tonsil size. To date, we have not identified, in previously published literature, any association between a high tongue position, reflected in FTP grade, and a concentric collapse pattern found on DISE.

Collapse	FTP I (n=23)	FTP II (n=48)	FTP III (n=95)	FTP IV (n=39)	P-value
PARTIAL	5 (21.7%)	3 (6.3%)	5 (5.3%)	2 (5.1%)	
COMPLETE					
Sum of sites					
I	2 (8.7%)	20 (41.7%)	29 (30.5%)	9 (23.1%)	
2	12 (52.2%)	16 (33.3%)	45 (47.4%)	21 (53.8%)	
≧3	4 (17.4)	9 (18.8%)	16 (16.8%)	7 (17.9%)	0.078
Sites					
Palatopharynx	7 (30.4%)	19 (39.6%)	46 (48.4%)	21 (53.8%)	
Hypopharynx	3 (13%)	6 (12.5%)	6 (6.3%)	2 (5.1%)	
Combined	8 (34.8%)	20 (41.7%)	38 (40%)	14 (35.9%)	0.261

Table 2 Partial Co	llapse and Sites of	Complete Collapse	e Among Different FTP Grades

**Notes:** Partial collapse: 50% to 75% lumen reduction; complete collapse: 75% to 100% lumen reduction. Sum of sites: I, one site showed complete collapse; 2, two sites showed complete collapse;  $\geq$  3, three or four sites showed complete collapse. Palatopharynx, velum and/or oropharynx sites; hypopharynx, tongue base and/or epiglottis sites; combined, palatopharynx and hypopharynx.

Abbreviations: FTP, Friedman tongue position; I, uvula and tonsils completely visible; II, visible uvula but not tonsils; III, visible soft palate, but not uvula; IV, visible hard palate only.

Site	Pattern	Degree	FTP I	FTP II (n=48)	FTP III	FTP IV (n=39)	<i>P</i> -value
			(n=23)		(n=95)		
			n (%)	n (%)	n (%)	n (%)	
V	AP	0	l (4.3%)	5(10.4%)	3(3.2%)	2(5.1%)	
		1 2	4(17.4%) 8(34.8%)	4(8.3%) 25(52.1%)	7(7.4%) 47(49.5%)	0(0%) 16(41%)	0.144
	L	0 I 2	0(0%) I (4.3%) 2(8.7%)	0(0%) 0(0%) 0(0%)	0(0%) 1(1.1%) 4(4.2%)	0(0%) 0(0%) 0(0%)	0.143
	с	0 1 2	0(0%) 3(13%) 4(17.4%)	0(0%) 3(6.3%) 11(22.9%)	0(0%) I(1.1%) 32(33.7%)	l (2.6%) l (2.6%) l 9(48.7%)	0.014
0	L	0 1 2	8(34.8%) 7(30.4%) 8(34.8%)	19(39.6%) 19(39.6%) 10(20.8%)	25(26.3%) 38(40%) 32(33.7%)	8(20.5%) 10(25.6%) 21(53.8%)	0.061
Т	AP	0 1 2	6(26.1%) 7(30.4%) 10(43.5%)	15(31.3%) 13(27.1%) 20(41.7%)	22(23.2%) 43(45.3%) 30(31.6%)	9(23.1%) 19(48.7%) 11(28.2%)	0.342
E	AP	0 1 2	16(69.6%) 0(0%) 6(26.1%)	33(68.8%) 2(4.2%) 11(22.9%)	61(64.2%) 9(9.5%) 24(25.3%)	26(66.7%) 6(15.4%) 6(15.4%)	
	L	0    2	0(0%) 0(0%) I (4.3%)	0(0%) 1(2.1%) 1(2.1%)	0(0%) 0(0%) 1(1.1%)	0(0%) I (2.6%) 0(0%)	0.337

Table 3 Collapse Pattern and Degree Among Different FTP Grades

Notes: 0, no collapse with 0% to 50% lumen reduction; I, partial collapse with 50% to 75% lumen reduction; 2, complete collapse with 75% to 100% lumen reduction. Abbreviations: FTP, Friedman tongue position; I, uvula and tonsils completely visible; II, visible uvula but not tonsils; III, visible soft palate, but not uvula; IV, visible hard palate only; V, velum; O, oropharynx; T, tongue base; E, epiglottis; AP, anterior-posterior; L, lateral; C, concentric.

In this study, the overall proportion of velum-CCC was 31.2%, consistent with that of large-sample studies that reported 31.5% and 37.3% in 1249 and 424 individuals with OSA, respectively.<sup>27,28</sup> This collapse pattern increased from 17.4% in FTP grade I to 48.7% in grade IV. In previous studies, an increased CCC at the retropalatal site was found

to be associated with obesity.<sup>27,29</sup> In this study, we found that a CCC increase was associated with a higher FTP grade, especially grade IV. Interactions between UA anatomy and muscle function may contribute to this result. In previous research, a high FTP grade was usually accompanied by a low hyoid position and incurred a higher

Table 4 Association Among FTP Grade IV and Characteristics of Complete Collapse Sites by DISE

Site	Pattern	Degree	Unadjusted OR (95% CI)	P-value	Adjusted OR (95% CI)	P-value
Velum	AP	2	1.3 (0.45–3.8)	0.626	1.5(0.49-4.59)	0.48
	L	2	0	1. The second se	0	1
	с	2	4.51 (1.3–15.71)	0.018	4.4 (1.19–16.23)	0.026
Oropharynx	L	2	2.19 (0.75–6.34)	0.149	2.04 (0.65–6.42)	0.223
Tongue base	AP	2	0.51 (0.17–1.5)	0.223	0.54 (0.18–1.64)	0.281
Epiglottis	AP+L	2	0.42 (0.12–1.44)	0.166	0.48 (0.14–1.71)	0.258

Notes: 2, complete collapse with 75% to 100% lumen reduction; OR adjusted for age, gender, body mass index, tonsil size grade with FTP I and tonsil grade 0 as reference. Abbreviations: FTP, Friedman tongue position; I, uvula and tonsils completely visible; II, visible uvula but not tonsils; III, visible soft palate, but not uvula; IV, visible hard palate only; DISE, drug-induced sleep endoscopy; AP, anterior-posterior; L, lateral; C, concentric. prevalence of CCC, as proven by sleep magnetic resonance imaging in severe OSA patients.<sup>19,30</sup> Moreover, the lateral muscles of the tongue and soft palate were thought to be coupled.<sup>31</sup> The circumference pattern may be explained by the increase in magnitude of the tongue drop, incurring pharyngeal occupation and central AP compression and posterior movement of the lateral margins of the soft palate, constituting a circumferential constriction.

As known, the presence of CCC was associated with clinical features in OSA patients, including higher respiratory disturbance index and AHI,<sup>32</sup> a more severe hypoxemia,<sup>33</sup> and susceptibility to non-positional OSA.<sup>28</sup> This collapse pattern can increase CPAP titration pressure requirement and rates of surgical failure,<sup>34,35</sup> as observed in UPPP and hypoglossal nerve stimulation (HGNS).<sup>36–38</sup> This association also explains why a high tongue position, especially FTP IV, is a predictor of severe OSA,<sup>39,40</sup> and causes significant reduction in AHI improvement after HGNS.<sup>41</sup> In this context, the identification of velum-CCC based on its association with FTP grade, as observed by a simple physical examination, has great clinical relevance in individuals with OSA.

Previous research demonstrated that a high tongue position indicated a more severe hypopharyngeal obstruction on DISE.<sup>42</sup> However, there was no statistically significant difference in this study, in which a complete tongue base collapse was identified in 30.6% of individuals with FTP grades III and IV, while Wang et al found an incidence of 32% in individuals with MMC classes III and IV.<sup>42</sup> In addition, we found no association between retroglossal collapse and tongue position on DISE, similar to the findings of den Herder et al.<sup>43</sup> Tongue base collapse has a more complex pathogenesis, related not only to tongue position and/or volume but also to muscle tone during sleep.<sup>44,45</sup>

There were some limitations to this study to consider. (i) Age was significantly different between groups, and the effect of aging on velum-CCC or AP collapse patterns is controversial.<sup>46,47</sup> Moreover, age was included in the logistic regression model to control for its influence. (ii) As a UA collapsibility evaluation method, DISE itself has some limitations, such as controversy about the reliability of observed obstructive sites,<sup>48</sup> that it might not improve surgical success rates,<sup>49</sup> and variable findings with prolonged observation time or different sedative drugs.<sup>44,50</sup> Dexmedetomidine was used for sedation, and differences in tongue base collapse related to propofol or midazolam use have already been identified. Nevertheless, the velum collapse pattern was similar in response to the three drugs,<sup>51</sup> enabling extension of the findings in this study to DISE with these drugs. (iii) In the sample, only OSA individuals intolerant to CPAP were evaluated, generating a selection bias, and not fully representing the general OSA population. (iv) The sample number in this study was relatively small, especially for the FTP IV group, probably leading to a low statistical power. More observation will be conducted in the future study.

## Conclusion

This study identified a positive correlation between a higher tongue position and retropalatal concentric collapse in OSA patients. FTP grade IV was an independent risk factor for CCC when controlling for age, gender, BMI, and tonsil size.

## **Ethics Approval**

All procedures performed in this study were in accordance with the ethical standards of the Institutional Review Board and Hospital Research Ethics Committee of Stanford Hospital and Clinics (Protocol 35,054). Patient informed consent was waived because of the retrospective nature of this research. The patients' data were maintained with confidentiality and in compliance with the Declaration of Helsinki.

## Disclosure

Dr Robson Capasso reports personal fees from and served at the advisory board for Invicta Medical, outside the submitted work. The authors report no other conflicts of interest in this work.

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