

Craniofacial and upper airway profile assessment in North Indian patients with obstructive sleep apnea

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ABSTRACT

Introduction: Upper airway imaging can often identify the anatomical risk factors for sleep apnea and provide sufficient insight into the pathophysiology of obstructive sleep apnea (OSA). **Materials and Methods:** We conducted a case-control, observational study at a tertiary care hospital in North India. All cases and controls underwent lateral cephalometry and magnetic resonance imaging (MRI) for craniofacial and upper airway evaluation. Only the cases had polysomnography testing for confirmation of OSA and assessing the severity of disease. **Results:** Forty cases and an equal number of matched controls were recruited. On X-ray cephalometry, it was observed that the cases had a significantly larger hyoid mandibular distance and soft palate length; and shorter mandibular length. The MRI cephalometric variables were significantly different, the soft palate length, tongue length, and submental fat were longer while the retropalatal and retroglossal distance was shorter amongst the cases. A statistically significant positive correlation was found between the cephalometric parameters and the indices of severity of OSA. An increased hyoid mandibular distance and soft palate length, and a decrease in the lower anterior facial height were found to be predictive of severe OSA (Apnea-Hypopnea Index \rightarrow 30/h). An increased hyoid mandibular distance, soft palate length, and the tongue length and a reduced mandibular length were found to be predictive of need for continuous positive airway pressure (CPAP) pressures of ≥ 15 cm H₂O. There were significant differences between the cephalometric parameters of the Indian OSA patients and patients from other ethnicities reported in the literature. **Conclusions:** OSA patients had a significantly smaller upper airway compared to age-, sex-, and body mass index-matched controls and cephalometric variables correlated with the indices of OSA severity. The cephalometric assessment was also predictive of severe OSA and the need for higher pressures of CPAP. This indicates the important role of upper airway anatomy in the pathogenesis of OSA.

KEY WORDS: Obstructive sleep apnea, magnetic resonance imaging, X-ray cephalometry

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INTRODUCTION

Obstructive sleep apnea (OSA) constitutes a spectrum of disorders of varying severity ranging from intermittent snoring at one end to obesity hypoventilation syndrome at the other end of the spectrum.^[1] OSA is increasingly being recognized as a disease with significant short- and

long-term cardiorespiratory complications. The prevalence rate of OSA in the middle-age population was reported to be 2%–4% in Caucasians.^[2] The overall prevalence of OSA in the adult population of Delhi was found to be 4.3% in a questionnaire-based study.^[3]

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Obesity has been frequently cited as a risk factor for OSA. However, OSA syndrome is known to occur even in nonobese patients where the other risk factors need to be ascertained. It is well recognized that OSA patients have an anatomically small upper airway with alterations in bony craniofacial structure and enlargement of surrounding soft-tissue structures;^[4] this spectrum of abnormalities acting synergistically to promote upper airway obstruction during sleep. The differences in the upper airway anatomy have been used to explain why nonobese patients can develop OSA.

Upper airway imaging is often not a part of routine evaluation in the diagnosis of OSA. Evaluation of the upper airway is usually done with the help of imaging methods such as X-ray lateral cephalometry, computed tomography (CT), magnetic resonance imaging (MRI), and nasopharyngoscopy. These different techniques are often complementary to each other; while X-ray allows only a two-dimensional detailed evaluation of the bony structures; for the three-dimensional reconstruction of the airway and soft-tissue structures (i.e., tongue, soft palate) techniques such as CT/MRI are better. Although these static imaging techniques are often not accurate in identifying the sites/sites of obstruction, they can identify the anatomic risk factors for sleep apnea and provide sufficient insight into the pathophysiology of OSA. Since individual patients have different patterns of upper airway narrowing, no single method for evaluating the obstruction appears to be complete in itself.^[5]

Multiple studies have documented differences in craniofacial features among the different races.^[6-8] Ethnic and racial differences in the upper airway dimensions need to be taken into consideration while comparing Indians with other ethnic groups. There is a paucity of data from India.^[9] We conducted this study to elucidate the differences in the upper airway anatomy between Indian OSA patients and controls; compare the cephalometric characteristics between the obese and nonobese OSA patients and also to determine the ethnic variations between Indian patients and other groups.

MATERIALS AND METHODS

We conducted a case-control observational study at a tertiary care hospital of a medical college in North India, after obtaining the institutional review board approval. Written, informed consent was obtained from the participants. The “cases” group comprised consecutive patients of either sex, 30–65 years of age, with clinical suspicion of OSA, attending the sleep clinic and willing to undergo polysomnography, arterial blood gas analysis, lateral cephalometry, and MRI. The “control” group comprised of patients visiting the hospital and undergoing an MRI for other reasons (for example undergoing MRI for evaluation of backache, headache) with no history suggestive of OSA and a negative response to the Berlin Questionnaire. The controls were matched for age, sex, and

body mass index (BMI) with the cases and were willing to undergo lateral cephalometry and MRI. Patients who had undergone any kind of maxillofacial or upper airway surgery or significant facial trauma in the past and pregnant females were excluded from the study.

Forty cases and an equal number of age-, sex-, and BMI-matched controls were recruited. All cases and controls were evaluated using predesigned pro forma for history taking and physical examination and underwent lateral cephalometry and MRI for craniofacial and upper airway evaluation. Only the “cases” had arterial blood gas analysis and polysomnography testing for confirmation of OSA and assessing the severity of disease.

Cephalometry

X-ray lateral cephalometry

Standardized lateral digital X-ray of head and neck was obtained in standing, natural head posture at end expiration, without swallowing, in centric occlusion and analyzed for hyoid mandibular distance, soft palate length, mandibular length, lower anterior facial height, and facial axis angle [Figure 1].^[7]

Magnetic resonance imaging

MRI of head, face, and neck was done using 1.5 Tesla Philips Achieva Scanner in supine position, while awake and both sagittal and transverse images were analyzed for soft palate length (craniocaudal), tongue length (craniocaudal), retropalatal and retroglossal oropharynx width (minimum), and submental fat thickness [Figures 2 and 3].^[10]

The aforementioned cephalometric and MRI parameters were selected because these parameters indicate the

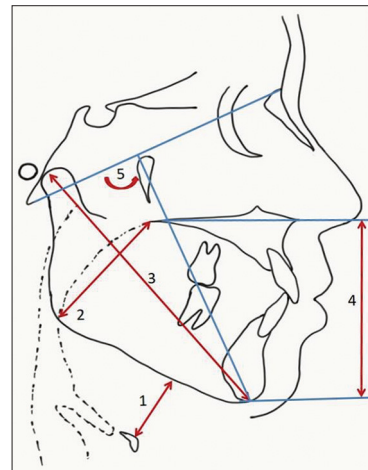


Figure 1: Measurements using lateral cephalometry. 1: Hyoid mandibular Distance: Perpendicular distance between hyoid bone and mandibular plane. 2: Soft palate length: Posterior nasal spine to inferior angle of the mandible. 3: Mandibular length: The distance between condylion to gnathion. 4: Lower anterior facial height: The distance between anterior nasal spine and menton measured perpendicular to the line passing through anterior nasal spine and the posterior nasal spine. 5: Facial axis angle: The angle formed by the basion-nasion and the plane from foramen rotundum to gnathion

size of the oropharynx and hypopharynx and are the most commonly implicated sites of obstruction in OSA patients.^[4-5,11]

Polysomnography study

Split-night polysomnography study (Alice 5 Diagnostic Sleep System, Respironics, USA) was conducted for the “cases” group in the sleep laboratory of the department and analyzed for Apnea-Hypopnea Index (AHI), Respiratory Disturbance Index, Arousal Index, and PAP requirement in OSAHS patients.

Statistical analysis

Hypotheses testing procedure was done with unpaired *t*-test with the level of significance set at 95% (alpha error – 5%, two-tailed test, and power – 80%). Sample size was calculated to detect a minimum difference of 2 mm in the retropalatal distance with a standard deviation of 3 mm.^[10]

Presence or absence of significant correlation between upper airway indices and severity of OSA was established using value of Pearson’s coefficient of correlation. Further data analysis was done using *t*-test for numerical data and Chi-square test for categorical data. Logistic regression analysis was done to estimate the odds ratio of predicting the needs for high CPAP (>15cm H2O). The odds ratio was adjusted for the BMI.

RESULTS

Forty cases and an equal number of age-, sex-, and BMI-matched controls were recruited. The baseline characteristics of both the groups are shown in Table 1. On physical examination, the neck circumference was larger and the incidence of retrognathia was found to be significantly higher in the OSA patients. A neck

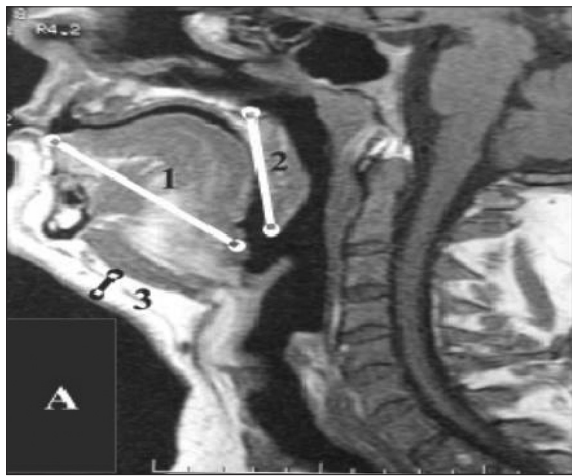


Figure 2: Magnetic resonance imaging sagittal section showing 1: Craniocaudal length of the tongue (maximum), 2: Craniocaudal soft palate length (maximum), 3: Submental fat thickness (maximum)

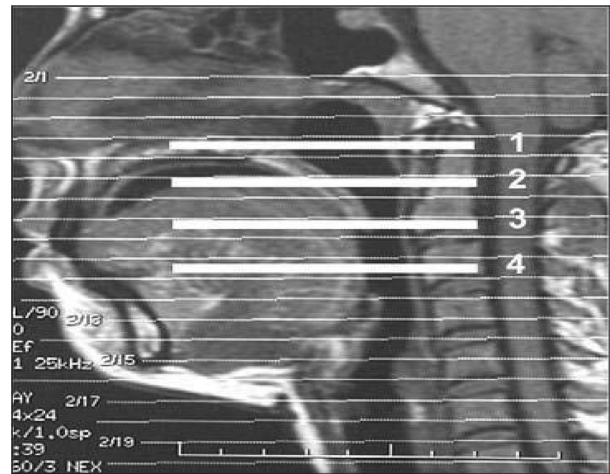


Figure 3: Magnetic resonance imaging sagittal section showing 1: Rhinopharynx (not measured), 2: High retropalatal oropharynx, 3: Low retropalatal oropharynx, retropalatal oropharynx (minimum) – measured between 2 and 3, 4: Retroglossal oropharynx (minimum)

Table 1: Baseline characteristics and physical examination findings of cases and controls

Parameter	Cases (n=40)	Controls (n=40)	P
Age (years)	48.80±7.68	48.45±8.09	0.843
Sex			
Males	31	31	1.000
Females	9	9	
BMI (kg/m ²)	29.96±2.66	29.82±1.96	0.784
Hypertension	20 (50)	4 (10)	<0.001
Diabetes	9 (22.5)	2 (5)	0.048
Epworth sleepiness scale	16.85±4.81	4.92±2.12	<0.001
Clinical examination			
Mean neck circumference (cm)	43.28±1.97	36.00±1.01	<0.001
Deviated nasal septum	1 (2.5)	0 (0)	1.000
Hypertrophied inferior nasal turbinate	2 (5)	0 (0)	0.494
Retrognathia	13 (32.5)	1 (2.5)	0.001
High-arched palate	7 (17.5)	1 (2.5)	0.057
Modified Mallampati classification			
Class 1	4 (10)	6 (15)	0.007
Class 2	9 (22.5)	20 (50)	
Class 3	21 (52.5)	14 (35)	
Class 4	6 (15)	0 (0)	

BMI: Body mass index

circumference of 40 cm or more was observed in all the cases and none of the controls. The case group comprised of more patients in the Modified Mallampati classification Class 3 and 4 than the control group. A higher Modified Mallampati Class indicated an overall smaller oropharynx with bulky tongue.

On comparing the X-ray cephalometric parameters between the cases and controls [Table 2], it was observed that the hyoid mandibular distance and the soft palate length were significantly larger among the cases and the mandibular length was significantly shorter in OSA patients. All the MRI cephalometric variables were significantly different between the two groups, the soft palate length, the tongue length, and

the submental fat were longer while the retropalatal and the retroglossal distance was shorter among patients with OSA. The length of the soft palate was measured using both lateral cephalometry as well as MRI and the measurements were consistent and comparable using both methods (correlation $r = 0.998$; $P < 0.001$). We also observed that patients had multiple cephalometric variables beyond the normal range, suggesting a reduction in the airway size at multiple levels. When we compared the cephalometric variables between the obese and nonobese OSA patients [Table 2], it was observed that the obese patients had a smaller airway dimension compared to nonobese patients though the amount of submental fat in both the groups was comparable.

Table 2: Comparing various cephalometric parameters between cases and controls and between the obese and the nonobese obstructive sleep apnea patient

Parameter	Mean±SD (95%CI)		P	Mean±SD (95%CI)		P
	Cases	Controls		Obese OSA	Nonobese OSA	
X-ray lateral cephalometry						
Hyoid mandibular distance (mm)	20.35±4.17 (19.02-21.68)	8.25±0.87 (7.97-8.53)	<0.001	21.90±3.31 (20.35-23.45)	18.80±4.43 (16.73-20.87)	0.017
Soft palate length (mm)	47.35±4.17 (46.02-48.68)	35.12±1.32 (34.70-35.55)	<0.001	48.80±3.19 (47.31-50.29)	45.90±4.59 (43.75-48.05)	0.026
Mandibular length (mm)	90.05±3.99 (88.77-91.33)	112.05±1.84 (111.46-112.64)	<0.001	88.70±4.32 (86.68-90.72)	91.40±3.20 (89.90-92.90)	0.031
Lower anterior facial height (mm)	73.78±5.52 (72.01-75.54)	74.68±5.24 (73.00-76.35)	0.457	72.60±5.88 (69.85-75.35)	74.95±5.01 (72.61-77.29)	0.182
Facial axis angle (°)	99.82±5.15 (98.18-101.47)	100.62±4.26 (99.26-101.99)	0.451	100.05±6.29 (97.10-103.00)	99.60±3.84 (97.80-101.40)	0.786
MRI variables						
Soft palate length (mm)	47.32±4.12 (46.01-48.64)	35.12±1.32 (34.70-35.55)	<0.001	48.70±3.15 (47.23-50.17)	45.95±4.58 (43.81-48.09)	0.033
Tongue length (mm)	74.18±4.74 (72.66-75.69)	71.50±3.72 (70.31-72.69)	0.006	75.65±4.17 (73.70-77.60)	72.70±4.92 (70.40-75.00)	0.048
Retropalatal distance (mm)	4.05±0.99 (3.73-4.37)	8.10±0.59 (7.91-8.29)	<0.001	4.10±1.21 (3.53-4.67)	4.00±0.73 (3.66-4.34)	0.753
Retroglossal distance (mm)	13.35±1.33 (12.92-13.78)	17.30±0.72 (17.07-17.53)	<0.001	13.15±1.59 (12.40-13.90)	13.55±0.99 (13.08-14.02)	0.349
Submental fat (mm)	14.12±1.68 (13.59-14.66)	9.15±0.95 (8.85-9.45)	<0.001	14.45±1.50 (13.75-15.15)	13.80±1.82 (12.95-14.65)	0.226

SD: Standard deviation, CI: Confidence interval, OSA: Obstructive sleep apnea, MRI: Magnetic resonance imaging

Table 3: Correlation of cephalometric indices with the neck circumference and the polysomnography findings

Upper airway parameter	Neck circumference (n=80)		RDI (n=40)		AHI (n=40)		AI (n=40)		CPAP pressure (n=40)	
	Correlation coefficient (r)	P	Correlation coefficient (r)	P	Correlation coefficient (r)	P	Correlation coefficient (r)	P	Correlation coefficient (r)	P
X-ray lateral cephalometry										
Hyoid mandibular distance	0.906	<0.001	0.825	<0.001	0.843	<0.001	0.420	0.007	0.909	<0.001
Soft palate length	0.910	<0.001	0.810	<0.001	0.823	<0.001	0.432	0.005	0.891	<0.001
Mandibular length	-0.915	<0.001	-0.590	<0.001	-0.597	<0.001	-0.420	0.007	-0.563	<0.001
Lower anterior facial height	-0.130	0.250	-0.180	0.265	-0.178	0.271	0.019	0.906	-0.146	0.369
Facial axis angle	-0.077	0.500	0.121	0.458	0.118	0.470	0.268	0.095	0.207	0.200
MRI variables										
Soft palate size	0.910	<0.001	0.800	<0.001	0.814	<0.001	0.432	0.005	0.895	<0.001
Tongue length	0.347	0.002	0.525	<0.001	0.528	<0.001	0.398	0.011	0.611	<0.001
Retropalatal distance	-0.865	<0.001	-0.246	0.125	-0.242	0.133	-0.310	0.052	-0.347	0.028
Retroglossal distance	-0.831	<0.001	-0.267	0.096	-0.269	0.093	-0.274	0.087	-0.255	0.112
Submental fat thickness	0.978	<0.001	0.496	0.001	0.508	<0.001	0.249	0.122	0.535	<0.001

RDI: Respiratory Disturbance Index, AHI: Apnea-Hypopnea Index, AI: Arousal index, CPAP: Continuous positive airway pressure, MRI: Magnetic resonance imaging

A statistically significant positive correlation was found between the cephalometric parameters and the indices of severity of OSA [Table 3]. Logistic regression analysis was done to determine the cephalometric variables significantly predictive of severe OSA (defined as an AHI of >30/h) [Table 4], it was seen that an increased hyoid mandibular distance was found to be predictive of severe OSA (odds ratio [OR] = 1.85; 95% confidence interval [CI] = 1.18, 2.89. $P = 0.007$). In addition, it was seen that an increased lower anterior facial height lowered the risk of severe OSA (OR = 0.86; 95% CI = 0.74, 0.99. $P = 0.045$). It was also observed that an increase of the soft palate length by 1 mm almost doubled the risk of having severe OSA. A similar analysis [Table 4] was done to look for parameters predictive of high continuous positive airway pressure (CPAP) (CPAP of ≥ 15 cmH₂O) requirement – an increased hyoid mandibular distance, soft palate length, and the tongue length and a reduced mandibular length were predictive of need for CPAP pressures of ≥ 15 cmH₂O.

The values of various anthropometric and upper airway indices obtained using lateral cephalometry and MRI in the OSA patients were compared with the corresponding values published in the literature to find any significant differences due to ethnic variations [Table 5]. On comparing the neck circumference and BMI with the other ethnicities, it was observed that the Indian patients had a greater BMI and larger neck circumference than the Far East Asians. Hyoid mandibular distance and mandibular length were significantly smaller in Indian OSA patients as compared to the Caucasians, suggesting a smaller hypopharynx in the Indian OSA patients as compared to the Caucasians. In comparison with the Japanese patients, mandibular length was significantly smaller in our OSA patients, whereas hyoid mandibular distance and the facial axis angle were significantly

greater in the Indians. In comparison with studies reported from Brazil, the Indian OSA patient had a significantly larger soft palate length.

DISCUSSION

In the present study, we demonstrated that OSA patients had a significantly smaller upper airway compared to age-, sex-, and BMI-matched controls and cephalometric variables correlated with the indices of OSA severity. The cephalometric assessment was also predictive of severe OSA and the need for higher pressures of CPAP. This indicates the important role of upper airway anatomy in the pathogenesis of OSA which is independent of obesity.

A high-arched palate, long uvula, tonsil enlargement, retrognathia, and obesity have been commonly reported in OSA patients.^[22] The current study identified a higher Mallampati score, retrognathia, and an increased neck circumference as risk factors for OSA. Ardelean *et al.*^[23] in a study from Europe had suggested a neck circumference cutoff of 41 cm, while in a study from Korea, Kang *et al.*^[24] determined the cutoff value for predicting OSA to be 34.5 cm. We observed that a neck circumference of 40 cm or more identified all the patients with OSA in our study population.

The hyoid mandibular distance was significantly increased, suggesting a lower placed hyoid bone in patients with OSA. Similar findings have been reported by Sforza *et al.*^[16] who suggested that the increased hyoid mandibular distance causes greater upper airway collapsibility. The hyoid bone position is believed to be crucial for pharyngeal patency and an imbalance between suprahyoid and infrahyoid muscles may influence the hyoid bone position.^[16] Skinner *et al.*^[25] who investigated the efficacy

Table 4: Logistic regression model for predicting severe obstructive sleep apnea and predicting the need for high continuous positive airway pressure (>15 cm H₂O)

Variable	Logistic regression model for predicting severe OSA			Logistic regression model for predicting need for high CPAP pressure (>15 cm H ₂ O)						
	OR	95% CI	P	Unadjusted OR			OR Adjusted for BMI			
				OR	95% CI	P	OR	95% CI	P	
Age	1.04	0.95-1.14	0.326	0.98	0.89-1.06	0.595	-	-	-	
BMI (kg/m ²)	1.15	0.89-1.49	0.294	1.58	1.13-2.22	0.008	-	-	-	
Neck circumference (cm)	1.47	0.96-2.23	0.074	1.41	0.96-2.06	0.078	-	-	-	
X-ray cephalometry										
Hyoid mandibular distance (mm)	1.85	1.18-2.89	0.007	6.11	1.69-22.11	0.006	5.44	1.49-19.87	0.010	
Soft palate length (mm)	1.92	1.22-3.00	0.004	2.89	1.26-6.65	0.012	2.62	1.11-6.17	0.028	
Mandibular length (mm)	0.87	0.73-1.04	0.122	0.68	0.54-0.87	0.003	0.75	0.58-0.98	0.038	
Lower anterior facial height (mm)	0.86	0.74-0.99	0.045	0.94	0.83-1.06	0.285	0.94	0.82-1.08	0.394	
Facial axis angle (°)	0.99	0.87-1.13	0.917	1.05	0.92-1.20	0.460	1.00	0.87-1.16	0.956	
MRI cephalometry										
Soft palate length (mm)	1.92	1.21-3.05	0.006	3.07	1.29-7.30	0.011	2.86	1.14-7.23	0.026	
Tongue length (mm)	1.08	0.94-1.24	0.281	1.24	1.03-1.51	0.022	1.15	0.94-1.41	0.159	
Retropalatal distance (mm)	0.97	0.51-1.87	0.933	0.31	0.11-0.86	0.024	0.29	0.09-0.94	0.039	
Retroglossal distance (mm)	0.89	0.54-1.49	0.665	0.83	0.51-1.37	0.472	0.88	0.52-1.48	0.634	
Submental fat (mm)	1.45	0.94-2.23	0.090	1.62	1.04-2.52	0.033	1.42	0.86-2.31	0.162	

BMI: Body mass index, MRI: Magnetic resonance imaging, OR: Odds ratio, CI: Confidence interval, CPAP: Continuous positive airway pressure, OSA: Obstructive sleep apnea

Table 5: Comparison of the anthropometric and cephalometric variables between different ethnic populations

	Indian OSA patients (n=40), Mean±SD		Caucasian OSA patients		Far East OSA patients		Brazilian OSA patients					
	Authors (References)	n	Mean±SD	P (compared to Indian patients)	Authors (References)	n	Mean±SD	P (compared to Indian patients)	Authors (Ref)	n	Mean±SD	P (compared to Indian patients)
Anthropometric parameters												
BMI	Lam <i>et al.</i> ^[12]	75	30±7	0.972	Lam <i>et al.</i> ^[12]	164	29±4	0.151	Borges <i>et al.</i> ^[13]	93	27.68±3.83	<0.001
	Liu <i>et al.</i> ^[14]	43	27.41±2.45	<0.001	Liu <i>et al.</i> ^[14]	30	26.98±2.49	<0.001				
	Li <i>et al.</i> ^[15]	293	30.7±5.9	0.435	Li <i>et al.</i> ^[15]	58	26.6±3.7	<0.001				
	Sforza <i>et al.</i> ^[16]	57	31.6±5.7	0.094	Kawaguchi <i>et al.</i> ^[17]	189	27.3±4.8	<0.001				
Neck Circumference	Lam <i>et al.</i> ^[12]	75	41±4	0.001	Lam <i>et al.</i> ^[12]	164	40±3	<0.001	Borges <i>et al.</i> ^[13]	93	38.56±3.92	<0.001
	Sforza <i>et al.</i> ^[16]	57	43.4±3.8	0.855	Kawaguchi <i>et al.</i> ^[17]	189	40.3±3.6	<0.001				
Cephalometric parameters												
Hyoid mandibular distance (mm)	Liu <i>et al.</i> ^[14]	43	26.27±6.27	<0.001	Liu <i>et al.</i> ^[14]	30	26.31±6.91	<0.001	Borges <i>et al.</i> ^[13]	93	19.21±8.22	0.407
	Li <i>et al.</i> ^[18]	50	26.4±7.1	<0.001	Li <i>et al.</i> ^[18]	50	18.7±6.5	0.167				
	Sforza <i>et al.</i> ^[16]	57	25.7±5.7	<0.001	Kikuchi <i>et al.</i> ^[19]	31	22.4±8.8	0.198				
Soft palate length (mm)	Liu <i>et al.</i> ^[14]	43	46.42±5.89	0.412	Liu <i>et al.</i> ^[14]	30	46.21±4.95	0.300	Borges <i>et al.</i> ^[13]	93	39.84±5.37	<0.001
	Sforza <i>et al.</i> ^[16]	57	47.2±4.9	0.875	Kikuchi <i>et al.</i> ^[19]	31	44.7±5.9	0.030				
Mandibular length (mm)	Liu <i>et al.</i> ^[14]	43	122.30±6.29	<0.001	Liu <i>et al.</i> ^[14]	30	118.56±7.46	<0.001				
	Sforza <i>et al.</i> ^[16]	57	123.0±6.3	<0.001								
Lower anterior facial height (mm)	Sforza <i>et al.</i> ^[16]	57	73.5±5.8	0.811								
Facial axis angle (°)												
Soft palate length (mm)	Rodenstein <i>et al.</i> ^[20]	10	47±7	0.851								
Retropalatal distance (mm)	Ciscar <i>et al.</i> ^[21]	17	39.1±4.9	<0.001								
	Rodenstein <i>et al.</i> ^[20]	10	4±1	0.887								
Retroglossal distance (mm)	Rodenstein <i>et al.</i> ^[20]	10	13±5	0.691								

BMI: Body mass index, OSA: Obstructive sleep apnea, SD: Standard deviation

of a titratable mandibular advancement splint (MAS) found that the baseline hyoid mandibular distance was the only cephalometric variable associated with a successful clinical outcome with the MAS.

The soft palate length was longer and the mandibular length was shorter in OSA patients than controls. This was further compounded by a larger tongue in such patients, causing a significant reduction in the size of the oropharynx. A large soft palate has been reported to be a common risk factor for OSA by Ciscar *et al.*^[21] and Sforza *et al.*^[16] Furthermore, pharyngeal occlusion likely occurs when the mandible is smaller or receded.^[26] Kim *et al.*^[27] reported that tongue volume and tongue fat were significantly enlarged in American patients with OSA when compared to obese controls. However, Okubo *et al.*^[28] from Japan did not find any significant difference in the tongue volume between OSA patients and controls. This might be due to ethnic differences. In the current study, we also observed the tongue length to be larger in the obese OSA patients than in the nonobese OSA patients though the submental fat was comparable between the groups.

The retropalatal and retroglossal distance in OSA patients was significantly lower as compared to controls indicating obstruction at the oropharynx level as an important pathogenetic factor. Our results were consistent with the findings of Ciscar *et al.*^[25] and Hora *et al.*^[29]

We also demonstrated that OSA patients had upper airway narrowing at multiple levels (retropalatal, retroglossal, shorter mandible, larger tongue, and lower hyoid bone position) with a combination present in most of the patients. Shellock *et al.*^[30] and Suto *et al.*^[31] had demonstrated multiple levels of occlusion and narrowing in the airway of OSA patients. The finding that the mixed type of pharyngeal obstruction was present in more than half of the patients has important clinical implications. Uvulopalatopharyngoplasty (UPPP) is the most common surgical procedure for patients with OSA. The success rate is related to the site of obstruction, with patients demonstrating retropalatal obstruction having better results than those with retroglossal obstruction.^[32,33] For patients in whom both retropalatal and retroglossal narrowing is seen, a surgery directed at advancing the tongue (e.g., geniohyoid advancement or maxillomandibular advancement) in addition to UPPP may be needed.

The current study showed that the lower anterior facial height and facial axis angle did not differ significantly between the cases and the control group. In contrast, Kikuchi *et al.*^[19] reported significantly lower facial axis angle and increased lower anterior facial height in the OSA patients. This, however, does not appear to be an important factor in Indian OSA patients indicative of important ethnic variations in the pathogenesis.

When we compared the obese and the nonobese patients, it was seen that the obese patients have a larger soft palate

and tongue, and associated anteroinferior positioning of the hyoid bone; suggesting a synergistic contribution of obesity and craniofacial factors to upper airway collapsibility. The crowding of the airway space through enlargement of the soft tissues could be the main catalyst for increased upper airway collapse in obese patients.

Ethnicity incorporates multiple factors such as obesity and craniofacial morphology, which will individually or in combination influence OSA.^[2] While Asian patients with OSA are generally less obese than their Caucasian counterparts, craniofacial abnormalities such as a low hyoid bone and repositioning of the maxilla or mandible are common predisposing factors for OSA in the Asian populations.^[34] In comparison, the Indian patients were found to be obese compared to Asians and to have smaller mandibular length compared to other ethnic groups. This suggests important differences in the bony facial structure and possible hints at a prominent role of MASs in the treatment of Indian OSA patients. Among soft-tissue parameters, tongue length was significantly smaller in Indian patients as compared to the Far East Asians, suggesting a less important role of tongue volume in the pathogenesis and treatment of OSA in Indian patients.

Strengths and limitations

The critical factors responsible for control of pharyngeal patency remain controversial. Two hypotheses have been proposed to explain the tendency of patients with OSA to collapse: aneural hypothesis implying reduced dilator muscle activity and an anatomic theory suggesting an anatomic narrowing of the upper airway. While we have assessed the anatomical factors in the current study, we also need to understand the contribution of neural factors; further studies using dynamic MRI and drug-induced sleep endoscopy will allow assessment of the dilator muscle activity.

Our study compared the craniofacial characteristics in OSA patients with BMI-matched controls, thus eliminating the impact of obesity on the upper airway profile. Use of MRI helped in better delineation of soft-tissue abnormalities and exact computerized measurements of several variables without exposing patients to unnecessary radiation. We also used an overnight PSG in all the patients and assessed the severity along with the CPAP pressures needed to treat the OSA. However, we could not do polysomnography to rule out OSA in the control group, although the sleep questionnaires were negative for the presence of OSA. The small sample size of the study limits the generalizability of the finding, and further studies are suggested.

Another limitation was that we compared the ethnic variations in upper airway indices of our OSA patients with the published literature, the effect of publication bias cannot be negated and the two groups might not be comparable.

CONCLUSIONS

The significant differences in the upper airway indices of OSA patients in comparison to BMI-matched controls signify the importance of anatomical features in the pathogenesis of OSA. The correlation of upper airway indices with the severity of OSA further demonstrates the impact of small changes in upper airway caliber on the severity of the disease. The identification of multiple sites of upper airway obstruction in majority of patients has important therapeutic implications. There may be differential contributions of craniofacial cephalometric dimensions and obesity to OSA between ethnic groups and particular ethnicities may be more vulnerable to changes in this relationship, such based on their anatomical substrate. More studies are needed to understand the complexity and interaction of OSA craniofacial phenotypes with obesity and also to assess the impact of ethnicity on these relationships.

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Conflicts of interest

There are no conflicts of interest.

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