Oropharyngeal examination as a predictor of obstructive sleep apnea
Pilot study of gag reflex and palatal reflex

Juliana Spelta Valbuza\textsuperscript{1,2}, Márcio Moysés de Oliveira\textsuperscript{1,2,3}, Cristiane Fiquene Conti\textsuperscript{1,2,3}, Lucila Bizari F. Prado\textsuperscript{1,2}, Luciane B.C. Carvalho\textsuperscript{1,2}, Gilmar Fernandes do Prado\textsuperscript{1,2}

ABSTRACT
Obstructive sleep apnea (OSA) has high prevalence and may cause serious comorbidities. The aim of this trial was to show if simple noninvasive methods such as gag reflex and palatal reflex are prospective multivariate assessments of predictor variables for OSA. \textbf{Method:} We evaluate gag reflex and palatal reflex, of fifty-five adult patients, and their subsequent overnight polysomnography. \textbf{Results:} Forty-one participants presented obstructive sleep apnea. The most relevant findings in our study were: [1] absence of gag reflex on patients with severe obstructive apnea (p=0.001); [2] absence of palatal reflex on moderate obstructive apnea patients (p=0.02). \textbf{Conclusion:} Gag reflex and palatal reflex, a simple noninvasive test regularly performed in a systematic neurological examination can disclose the impact of the local neurogenic injury associated to snoring and/or obstructive sleep apnea syndrome. \textbf{Key words:} obstructive sleep apnea, OSA, gag reflex, palatal reflex.

O avaliação orofaríngea como preditor da apneia obstrutiva do sono: estudo piloto dos reflexos nauseoso e palatal

RESUMO
A síndrome da apneia obstrutiva do sono (SAOS) possui alta prevalência e pode causar sérias comorbidades. O objetivo deste estudo foi mostrar se métodos não invasivos como os reflexos nauseoso e palatal podem ser avaliações prospectivas multivariadas preditoras para SAOS. \textbf{Método:} Avaliamos os reflexos palatal e nauseoso em 55 pacientes adultos, com exame polissonográfico subsequente. \textbf{Resultados:} 41 pacientes apresentaram SAOS. Os achados mais relevantes em nosso estudo foram: [1] ausência do reflexo nauseoso em pacientes com SAOS grave (p=0.001); [2] ausência do reflexo palatal em pacientes com SAOS moderada (p=0.02). \textbf{Conclusão:} Os reflexos nauseoso e palatal, um simples exame não invasivo, aplicado em uma avaliação neurológica rotineira, pode revelar o impacto de lesões neurogênicas locais associadas ao ronco e/ou a SAOS. \textbf{Palavras-Chave:} apneia obstrutiva do sono, SAOS, reflexo palatal, reflexo nauseoso.

Obstructive sleep apnea (OSA) is a common disorder characterized by recurrent episodes of upper airway obstruction during sleep. The upper airway anatomy and neuromuscular functions play a crucial role in the pathophysiology of OSA\textsuperscript{1-5}. Pathogenesis of airway obstruction in patients with OSA remains incompletely understood\textsuperscript{6}. The primary defect is probably an anatomically small or collapsible pharyngeal airway, in combination with a sleep-induced fall in upper airway muscle activity\textsuperscript{2}.

As mentioned in the study of Stuck and Maurer\textsuperscript{7}, airway anatomic variables related to OSA have been incorporated into complex models involving detailed
physical or radiographic measurements. Researchers and clinicians used different techniques to reveal potential differences in upper airway anatomy to better understand the pathophysiology of the disease but also to improve patient management and treatment.

There is evidence for impairment of upper airway mucosal sensory function in the oropharynx of patients with OSA. The pathophysiological mechanism is believed to be local neurogenic lesions in the oropharynx caused by the low-frequency vibration of habitual snoring in OSA.

Part of neurological examination is based on stimulus-response approach and the palate and pharynx should routinely be tested in the physical exam, but we assume a normal function of these structures when patients do not complain of dysphagia or choking, and we also rarely test gag reflex (GR) and palatal reflex (PR) in our daily clinics. The absence of GR and PR in the context of the neurological examination were not fully investigated in the scenario of obstructive sleep apnea.

The soft tissue trauma caused by vibration could develop secondary mechanical or inflammatory-related local neuropathy, what has been showed by studies using complex invasive techniques to assess the upper-airway sensation in OSA, such as endoscopic sensory tests, biopsies, and electrical stimulation.

We hypothesize that patients with obstructive sleep apnea have gag and palatal reflexes impaired, hence the aim of this study is to verify if the well known neurological physical examination, including GR, and PR are predictive of pharynx impairment in patients with obstructive sleep apnea syndrome. We justify our study because patient’s evaluation already include those approach, the procedures require no special equipment or skills, are simple to learn, non-invasive, and evidently low costing.

**METHOD**

**Participants**

We enrolled 55 consecutive obstructive sleep apnea and control participants from 2009 to 2010 from the Neuro-Sono Sleep Center and São Paulo Hospital Sleep Laboratory, Department of Neurology, Federal University of São Paulo (UNIFESP). The study protocol was approved by the local ethics committee and all participants signed a consent form.

**Patients**

Forty-one consecutive patients referred to the Neuro-Sono Sleep Center, Department of Neurology, complaining of snoring and daytime sleepiness were examined clinically and had a standard polysomnography (PSG) done. There were 22 male and 19 female, age from 27 to 76 years (55.2±14.2), and body mass index (BMI) from 18.5 to 46.6 Kg/m² (27.9±15.8). The characteristics of the forty-one patients are presented in Table 1.

**Controls**

The controlled group included 14 volunteers without snoring and/or sleep apnea, confirmed by polysomnography, randomly recruited from a pool of patients with normal PSG referred to our sleep center mostly for insomnia, parasomnia or periodic limb movements of sleep (PLMS). We matched them to the patients group according to the BMI, age, and gender. There were 7 male and 7 female, age from 17 to 78 years (42.6±14.0), and mean body mass index from 18.3 to 33.0 Kg/m² (24.2±5.3). The characteristics of the fourteen volunteers are presented in Table 1.

**Sleep study**

All patients had an overnight polysomnography including standard electroencephalographic leads, bilateral electrooculogram, chin and tibialis electromyograms, airflow via nasal pressure cannula and thermistor, thoracoabdominal movements via piezoelectric belt, body position via position sensor, and arterial oxyhemoglobin saturation via finger pulse oxymetry. All signals were acquired on a digital data-management system Neurotec model, EQSA-400, Itajuba, MG, Brazil. Studies were scored manually by trained, experienced clinical neurophysiologist blind for the purpose of this study. Sleep-wake state was defined according to standard criteria. Obstructive sleep apnea episodes were defined as cessation of airflow lasting at least 10 seconds with persistent respiratory effort, and hypopneas as episodes of reduction in airflow or inspiratory flow limitation on the nasal cannula pressure signal lasting more than 10 seconds with an associated desaturation of at least 3% or arousal defined according to American Sleep Disorders Association criteria. We considered mild OSAS when apnea-hypopnea index (AMI) was between 5 and 15, moderate when between 15 and 30, and severe when above 30/h.

**Table 1. Characteristics of participants.**

<table>
<thead>
<tr>
<th>N</th>
<th>Gender</th>
<th>Age (mean±SD)</th>
<th>BMI (mean±SD)</th>
<th>AHI (N%)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Group Control</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>14</td>
<td>07♂</td>
<td>42.6±14.0</td>
<td>24.2±5.3</td>
<td>Normal: 14 (100%)</td>
</tr>
<tr>
<td>07♀</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Group Patients</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>41</td>
<td>22♂</td>
<td>55.2±14.2</td>
<td>27.9±15.8</td>
<td>Mild: 19 (46.4%)</td>
</tr>
<tr>
<td>19♀</td>
<td></td>
<td></td>
<td></td>
<td>Moderate: 6 (14.6%)</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td></td>
<td>Severe: 16 (39%)</td>
</tr>
</tbody>
</table>

N: number of participants; SD: standard deviation; AHI: apnea-hypopnea index.
Palatal reflex

The palatal reflex was obtained during the physical examination of each patient. For all patients, the assessment of scores was done or directly supervised by the same physician. The reflex was assessed by asking the patient to open his/her mouth as wide as possible, while protruding the tongue as far as possible. The patient was instructed to not emit sounds during the assessment. A wooden spatula was placed in contact with the mucosa of right or left anterior pillar of the soft palate stimulating the contraction of soft palate (palatoglossus muscle, Fig 1), in case of Mallampati IV, when necessary we depressed the tongue with another wooden spatula to better see the anterior pillar of the soft palate. Palatal reflex was considered present when the soft palate moves upward or backward. When the reflex was to be absent, the stimulus was repeated after one minute in the contralateral area, to confirm the absence of reflection or not.

Gag (nauseous) reflex

To assess the gag reflex we adopted the same condition above described and we touched the wooden spatula on the mucosa of the posterior wall of the pharynx evoking the gag reflex (Fig 2), in case of Mallampati IV, when necessary we depressed the tongue with another wooden spatula to better see the posterior wall of the pharynx. Gag reflex was considered present when the pharyngeal wall contracted associated or not to soft palate contraction. When the reflex was to be absent, the stimulus was repeated after one minute in the contralateral area, to confirm the absence of reflection or not.

Statistical analysis

We used the Chi-square test and Fisher test to analyze the presence or absence of palatal reflex and gag reflex, comparing patients with mild, moderate, and severe OSAS to control group. We considered a p-value <0.05 for statistical significance.

RESULTS

Demographic data

We studied 55 individuals with suspected sleep disorders and who had been referred to the São Paulo Hospital Sleep Laboratory, 29 male and 26 female participants, with age from 17 to 78 years old, and body mass index from 18.3 to 46.6 Kg/m². Overall, forty-one of fifty-five individuals (74.5%) presented obstructive sleep apnea (19 mild, 06 moderate and 16 severe).

Palatal reflex

Overall the palatal reflex was absent on 12 participants (n=55; 21.8%). Palatal reflex was absent just in one patient of the control group (n=14; 7%); in 2 patients from mild group (n=19; 10.5%); in 4 patients from moderate group (n=6; 66.7%); and 5 patients from severe group (n=16; 31%). Compared to control group the absence of palatal reflex was significantly higher only in the moderate OSAS patients (66.7%; p=0.02) (Table 2).

Gag reflex

The gag reflex was absent on 12 participants (n=55; 21.8%) and none of the control group presented absence of gag reflex (n=14; 0%). The gag reflex was absent just

<table>
<thead>
<tr>
<th>Group</th>
<th>Reflex</th>
<th>N (reflex absent)</th>
<th>p value (compared to control)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Control</td>
<td>Palatal</td>
<td>1</td>
<td>-</td>
</tr>
<tr>
<td>N=14</td>
<td>AHI (N%)</td>
<td>Gag</td>
<td>0 (zero)</td>
</tr>
</tbody>
</table>
| Normal: 14 (100%) | Mild: 0 (0%) | Severe: 0 (0%) | Palatal Reflex was absent just in one patient of the control group (n=14; 7%); in 2 patients from mild group (n=19; 10.5%); in 4 patients from moderate group (n=6; 66.7%); and 5 patients from severe group (n=16; 31%). Compared to control group the absence of palatal reflex was significantly higher only in the moderate OSAS patients (66.7%; p=0.02) (Table 2).

Gag reflex

The gag reflex was absent on 12 participants (n=55; 21.8%) and none of the control group presented absence of gag reflex (n=14; 0%). The gag reflex was absent just
in one patient from mild group (n=19; 5.2%); one from moderate group (n=6; 16.6%); and ten patients from severe group (n=16; 62.5%). Compared to control group the absence of gag reflex was significantly higher only in severe OSAS patients (62.5%; p=0.001) (Table 2).

**DISCUSSION**

Upper airway (UA) tissues lesions on OSA patients are associated with a slowing of impulse conduction, these sensory impairment is caused by axonal degeneration and segmental demyelization in afferent neurons. There is morphologic evidence to support the presence of a sensory neuropathy in the UA in obstructive sleep-disordered breathing.

Several authors provided evidence for local neurogenic lesions in OSA. Most of these authors hypothesized that snoring was responsible for the histologic alterations reported.

These observations have led to the suggestion that the progression from mild snoring to heavy habitual snoring and then OSA may represent a progressive local neuropathy.

Overall we believe our findings are consistent with the UA sensory impairment developing as a consequence of snoring and while remaining partially reversible, representing a largely permanent injury, easily demonstrated by the physical examination of PR and GR, because our data showed that the absence of these reflexes, as defined in the study of Davies and Kidd, were more significant when associated to the severity of OSA.

Although, some inconsistent data such as PR, that showed a no significant result with severe OSA group, but with significant result with moderate OSA group, could be analyzed as a bias because of the small sample of participants, and the unknown illness time by the authors.

The UA sensory impairment could potentially represent a defect in the afferent limb of such protective reflexes, in concordance with our findings.

The independent association between Gag reflex, Palatal reflex and the presence and severity of OSA suggests that these scoring systems will have practical value in clinical settings and in prospective studies of sleep-disordered breathing.

We remain cautious in asserting that the use of PR or GR as a simple diagnostic test may be sufficient to predict OSAS, as demonstrated in other tests (questionnaires for OSA, outpatient recording). But we hope that future studies with larger numbers of participants, become an opportunity to be a predictor test for OSAS, even as it would be an important tool, easy to perform, which could prioritize patients for polysomnography, and important consideration given the large backlog of patients awaiting assessment for OSA.

We concluded that gag and palatal reflex were altered on OSA patients, suggesting them as predictor factors as well. Patients with OSA gradually have neurogenic injuries caused by trauma of snore, contributing for the instability of upper airway, proved by complex tests, such as sensory measures by endoscopic methods, and biopsies.

Our findings suggest that a simple noninvasive test like GR and PR physical examination, are associated to OSAS. We strongly encourage further trials to corroborate more evidence about PR and GR as predictor factors for OSA.

**REFERENCES**