# Treatment options for obstructive sleep apnea

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#### **Abstract**

Purpose of review: Obstructive sleep apnea (OSA) is a global problem with implications for general health and quality of life, and is often encountered in patients with neurologic disease. This review outlines treatment modalities to consider for management of OSA in patients with neurologic disease. Recent findings: New advances in positive airway pressure (PAP) devices, oral appliances, and surgical interventions offer a wide range of treatment options for patients with OSA. Summary: PAP therapy remains the gold standard treatment for OSA. Other treatment modalities may be considered for OSA patients who decline or cannot tolerate PAP therapy. Some OSA patients may benefit from multimodal treatment. Neurol Clin Pract 2017;7:77-85



atients with multiple sclerosis, epilepsy, neurodegenerative disease, and other categories of neurologic disease often look to the treating neurologist for evaluation and management of fatigue and tiredness. While fatigue may be a component of the underlying neurologic condition or medication side effect, it may also reflect comorbid obstructive sleep apnea (OSA). This brings recognition and treatment of OSA into the arena of the general neurologist, who often seeks to improve patients' daytime functioning without the addition of multiple medications. Having a working knowledge of OSA and available treatment options allows general neurologists to counsel patients about how identification and appropriate treatment of OSA may help improve daytime fatigue and reduce risk of comorbidities associated with untreated OSA.

OSA affects 9%–26% of the middle-aged population¹ and is characterized by ≥15 obstructive events per hour of sleep or ≥5 obstructive events per hour of sleep and the presence of any related symptoms (snoring, witnessed apnea, nocturnal gasping or choking, nonrestorative sleep, excessive daytime sleepiness).2 OSA incidence in patients with certain neurologic conditions, such as stroke or medication-refractory epilepsy, is higher than in the general population.<sup>3,4</sup> Untreated OSA can lead to daytime sleepiness, decreased productivity, increased motor vehicle accidents, and worsening hypertension, atrial fibrillation, and stroke.<sup>5,6</sup> Appropriate evaluation and treatment of OSA may yield symptomatic benefit for patients with comorbid neurologic disease.

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# CPAP is considered the gold standard treatment for OSA because its use can improve sleep-related symptoms and quality of life.

The table summarizes the OSA treatment options that are discussed in detail.

### What treatment should I consider first for my OSA patients?

Continuous positive airway pressure (CPAP) CPAP is considered the gold standard treatment for OSA<sup>7</sup> because its use can improve sleep-related symptoms and quality of life. CPAP acts as a pneumatic splint that stabilizes the upper airway with constant positive pressure via a mask interface. CPAP has been shown to significantly decrease the apnea-hypopnea index (AHI), an average that represents the combined number of apneas and hypopneas that occur per hour of sleep, particularly in patients with severe disease (defined as an AHI  $\geq$ 30/h).\(^1 CPAP users also show improvements in objective and subjective sleepiness compared to controls.\(^8

Despite these benefits, nightly CPAP use remains problematic for some patients. Difficulty with CPAP may stem from mask discomfort or claustrophobia, pressure intolerance, lifestyle or social considerations, or a combination of these factors.

CPAP: Consider this gold standard treatment modality as first-line for all OSA patients.

### What if my OSA patients have trouble tolerating the CPAP setting?

Individuals with pressure intolerance may experience dryness or irritation of nasal and pharyngeal membranes, nasal congestion, or eye irritation from air leakage with CPAP use. Use of heated humidification<sup>10</sup> or different mask types may facilitate comfort and increased CPAP use. Short-term reduction of the pressure setting<sup>10</sup> may help the patient become acclimated to

### Treatment options for obstructive sleep apnea Positive airway pressure (PAP) Continuous PAP (CPAP) Bi-level PAP +/- Backup rate +/- Average volume assured pressure support Autotitrating PAP (may be CPAP or bi-level PAP) Oral appliances Tongue-retaining devices Mandibular advancement devices Surgical Phase I (nasal, palatal, tongue) Phase II (maxillomandibular advancement) Hypoglossal nerve stimulation Adjunct Weight loss Positional therapy Nasal expiratory PAP Noninvasive oral pressure therapy

CPAP, with the ultimate goal to eventually resume the prescribed pressure. Flexible pressure at end-expiration may also increase patient comfort with the pressure.

Other influences on adherence include apprehension about how CPAP will affect appearance or interfere with sexual functioning, the patient's and bed partner's perceived benefit, if the patient sleeps alone, socioeconomic status, and anxiety or depression. Open discussions can help mitigate negative perceptions of CPAP therapy and work towards nightly CPAP use.

*Bi-level positive airway pressure (PAP)* Bi-level PAP was originally developed to administer varying pressures between the inspiratory and expiratory cycle, one pressure for inhalation and the second for exhalation. This may facilitate tolerance of higher pressures via easier exhalation. During the inspiratory cycle, the greater level of pressure combats the inspiratory flow limitation in the upper airway.<sup>11</sup> This yields a greater tidal volume and unloading of the respiratory muscles compared to CPAP and essentially treats nonobstructive sleep-related hypoventilation.<sup>11</sup> Some bi-level PAP devices have additional capabilities, including backup breath delivery in a spontaneous or timed mode.

Bi-level PAP with average volume-assured pressure support adapts pressure support to guarantee a predetermined tidal volume.<sup>12</sup> This noninvasive ventilation modality may be considered for patients with obesity hypoventilation, chronic obstructive pulmonary disorder, or alveolar hypoventilation seen in neuromuscular disorders such as amyotrophic lateral sclerosis (ALS), spinal muscular atrophy, myotonic dystrophy, or cervical spinal cord injury.<sup>12,13</sup> In one study comparing patients with little or no bulbar involvement ALS, the use of a noninvasive ventilator had been shown to increase median survival in the noninvasive ventilation group for 48 days longer for a total of 219 days vs 171 days in the standard care group.<sup>14</sup>

Bi-level PAP: Consider for OSA patients with intolerance to CPAP or those who have hypoventilation in the setting of obesity or neuromuscular disease.

Autotitrating PAP (APAP) APAP is set to a pressure range and the delivered pressure varies depending upon a device-specific algorithm. Both CPAP and bi-level PAP devices are available in autotitrating modes. Generally the device defines normal airflow for the patient, and delivers additional pressure when the flow curve is restricted. APAP devices may be useful for patients whose titration-recommended setting was not suitable due to the inherent limited sampling of a single night or half-night titration study. APAP therapy may also be considered for patients in whom large weight fluctuations are anticipated, such as pregnant patients or those undergoing bariatric surgery. Contraindications to APAP include congestive heart failure, lung disease such as chronic obstructive pulmonary disease, obesity hypoventilation syndrome, lack of snoring, or central sleep apnea.

APAP: Can consider for OSA patients who do not have clear contraindications to APAP use.

### Can my OSA patients use an oral appliance for treatment?

*Oral appliance therapy for OSA* Oral appliances can benefit carefully selected OSA patients who prefer not to use CPAP or have comprehensive surgeries. Oral appliances improve upper airway patency during sleep by enlarging the upper airway or by decreasing upper airway collapsibility. <sup>16</sup> Oral appliances can be categorized as tongue retaining devices and mandibular advancement devices (MADs), with the latter being more commonly used. MADs cover the upper and lower teeth and hold the mandible in a forward position with respect to the resting position and can be customizable or over-the-counter. <sup>16</sup> Mandibular protrusion advances the tongue position and subsequently increases oropharyngeal volume. <sup>17</sup>

### Is my OSA patient a candidate for a MAD?

MADs are indicated for patients with mild to moderate OSA or those with severe OSA who are unwilling or unable to use CPAP, have good dentition, and have a body mass index (BMI) below 30 kg/m<sup>2</sup>.<sup>2</sup> A dentist trained in dental sleep medicine examines the mandible, bite, dentition, and range of motion to identify patients with temporomandibular joint

dysfunction<sup>6</sup> that may worsen with MAD use. Accredited sleep laboratories may also offer polysomnographic evaluation with a remotely controlled mandibular positioner to help predict an effective mandibular protrusion position. This technology is currently available for clinical use at sleep laboratories in many geographic regions.<sup>18</sup>

In patients with mild to moderate OSA, a >50% reduction in AHI to <5/h occurred in 42.8% of patients using an oral appliance and 73.2% of patients using CPAP. <sup>19,20</sup> Odds of achieving AHI <5/h and <10/h was respectively 49 times greater and 89 times greater with the oral appliance-treated group compared to the control untreated group. <sup>19,20</sup> Compared to noncustom MADs, custom MADs have been shown to achieve greater reduction in AHI. <sup>19</sup>

Adherence to oral appliance therapy is difficult to track objectively and thus compare directly with rates of CPAP adherence. Thermosensors can be embedded into oral appliances to monitor device usage,<sup>21</sup> but presence of these sensors is not ubiquitous. Overall, the absolute difference between the mean subjective adherence rates for oral appliance users was 0.70 more hours per night than the objective adherence rates among CPAP users.<sup>19,20</sup>

MADs: Consider for patients with mild to moderate OSA, or those with severe OSA who have chosen not to use or are unable to use CPAP.

### What surgical treatment options are available to my OSA patients?

Because OSA relates to the structure and tone of the upper airway, upper airways surgeries are aimed at reducing the degree of obstruction in the nose, oropharynx, or hypopharynx. Anatomical features that may predispose to OSA include retroposition of the maxilla or mandible, enlarged pharyngeal fat pads, enlarged soft palate, enlarged tongue, narrow posterior airway space, and upper airway hypotonia.

Phase I surgeries include nasal surgeries, palatal surgeries, and tongue-based surgeries that may initially alleviate and treat OSA. Phase II surgery, known as maxillomandibular advancement, is generally reserved for patients with major OSA due to velo-orohypopharyngeal airway obstruction that could not be resolved from phase I surgeries.<sup>22-24</sup>

Surgical success is frequently defined when the individual achieves a greater than 50% reduction of the AHI or an AHI of ≤20/h.<sup>25,26</sup> However, this oft-used definition of success for OSA surgeries may not represent a clinical "cure," so careful attention must be paid to endpoints used in studies of surgical treatments for OSA.<sup>27</sup> Determination of surgical candidacy on a case-by-case basis, with attention to severity of OSA, presence/absence of obesity, and presence/absence of craniofacial features amenable to surgery, can help increase the likelihood of surgical success. Continuous weight management and optimal treatment of medical comorbidities is imperative to reduce postoperative worsening or recurrence of OSA.

# What are the initial (phase I) surgeries to consider for my OSA patients?

In general, phase I surgeries should be considered for patients with mild to moderate OSA who are nonobese.

**Adenotonsillectomy** Adenotonsillar hypertrophy has become the most common cause of pediatric OSA,<sup>28</sup> for which adenotonsillectomy may be clinically indicated as first-line treatment in certain children.<sup>29</sup> Palatine tonsil hypertrophy rarely causes adult OSA<sup>30</sup> and thus isolated adenotonsillectomy does not yield much improvement in OSA in adults.

Adenotonsillectomy: Can consider as first-line for appropriate pediatric OSA patients but is rarely used as a sole treatment for adult OSA patients.

*Nasal surgeries* Nasal obstruction due to turbinate hypertrophy or septal deviation may compound snoring and OSA.<sup>31</sup> Intranasal pathology increases upper airway resistance with subsequent collapse, which leads to hypopneas.<sup>31</sup> Surgical techniques such as septoplasty with turbinate cautery and removal of concha bullosa can decrease airway resistance, which can treat OSA in some and can facilitate CPAP comfort for others.<sup>31</sup>

Nasal surgery: May cure some patients of their OSA; may allow others to try a smaller mask with a lower CPAP setting.

Palatal surgeries Uvulopalatopharyngoplasty (UPPP) was developed to open a narrow or collapsed retropalatal area. Friedman et al.<sup>32</sup> developed a staging system that incorporates palate position and BMI to estimate the rate of postsurgical success. Compared to the Mallampati classification, which assess patency of the oral airway with the tongue protruded, the Friedman tongue position is assessed with the tongue in a natural, neutral position. The essence of the Friedman staging system considers patients with a patent oral airway and BMI <40 kg/m² as stage I; patients with a moderately crowded oral airway and BMI <40 kg/m² (any patency of oral airway) as stage III. Studies suggest that stages I, II, and III patients who underwent UPPP had respective success rates of 80.6%, 37.9%, and 8.1%.<sup>32</sup> Moderate to severe postoperative OSA<sup>25</sup> still requires treatment, either with PAP therapy or an alternative such as an oral appliance.

Other palatal procedures include palatal z-plasty and pillar procedure. The pillar procedure reduces snoring by slightly stiffening the soft palate to mitigate its compliance and was found to lower AHI from  $16.5 \pm 4.4$ /h to  $11.2 \pm 10.2$ /h in patients with mild to moderate OSA.<sup>33</sup> Palatal z-plasty, developed for post-tonsillectomy patients who continued to have air obstruction issues and were not candidates for an UPPP,<sup>34</sup> has been shown to reduce snoring in patients with Friedman palate position II and III and may also lower the AHI. In a prospective study comparing UPPP and palatal z-plasty, in which surgical success was defined as a minimum 50% reduction in AHI and postoperative AHI less than 20/h, surgical success occurred in 28% of UPPP patients with a preoperative AHI of  $33.4 \pm 13.9^{34}$  and in 68% of those who underwent palatal z-plasty with a preoperative AHI of  $41.8 \pm 26.4$ .<sup>34</sup>

Palatal surgeries: Patients with more a patent oral airway, nonsevere OSA, and BMI < 40 kg/m<sup>2</sup> may have better surgical outcomes.

**Tongue-based surgeries** Genioglossus advancement with hyoid suspension for treatment of OSA met the surgical criteria for success (defined as a minimum 50% reduction in AHI and a postoperative AHI less than 20/h) for 70% of the participants involved in the study.<sup>35</sup> Genioglossus advancement with hyoid suspension is often an adjunctive intervention with UPPP. Outcomes involving multiple oral-pharyngeal surgical interventions with UPPP showed an overall success rate of 76%.<sup>32,36</sup>

Transoral robotic surgery, developed to address tongue base and supraglottic obstruction, has been performed as a stand-alone procedure or as part of a multilevel pharyngeal, palatal, or nasal surgery. One study was able to demonstrate 84.3% of patients to have improvement in their preoperative vs postoperative AHI (42.7 vs 22.2, respectively). In this study, 51.2% of patients met the surgical criteria for success, and 14% met the criteria for cure as defined as AHI <5/h. Individuals with a BMI <30 kg/m² had a 15.3% cure rate, compared to 11.1% cure rate in those with a BMI >30 kg/m².

**Multilevel surgeries** Multilevel surgeries for OSA address multiple areas of obstruction along the upper airway. These may include nasal, palatal, and tongue surgeries,<sup>39</sup> while other articles have described only palatal and tongue-based surgeries.<sup>25</sup> Multilevel surgeries can be performed simultaneously or in a phased sequence, which may include phase I and phase II procedures.

Tongue-based surgeries: May be performed in isolation or as part of multilevel surgery; patients with  $BMI < 30 \text{ kg/m}^2$  may have better surgical outcomes.

# What are the subsequent (phase II) surgeries to consider for my OSA patients?

*Maxillomandibular advancement (MMA)* MMA enlarges the velo-orohypopharyngeal airway via advancement of anterior pharyngeal tissues attached to the mandible. This procedure is sometimes reserved for patients with major maxillomandibular deficiency. For the best outcome, surgical candidates should have a BMI of  $\leq 30 \text{ kg/m}^2$ , cephalometric posterior airway space  $\leq 9 \text{ mm}$  defining hypopharyngeal narrowing, velo-orohypopharyngeal narrowing, and potential retrognathia. Typically seen as a final approach in stepwise phase surgery, and deemed phase II, MMA alone has been shown

## Patients with OSA and comorbid obesity should be counselled on long-term weight management.

to be as effective as UPPP and MMA combined.<sup>41</sup> In a review of 234 individuals, 90% of whom were men with a mean BMI of 29.1 kg/m<sup>2</sup> and preoperative mean AHI of 54.4/h, MMA led to 87% reduction in AHI of with a mean postoperative AHI of 7.7/h.<sup>22-24</sup> Postsurgical facial aesthetics is generally discussed with the patients prior to the procedure. The maxillomandibular complex is advanced between 10 and 12 mm routinely in OSA patients, causing excessive maxillomandibular protrusion,<sup>42</sup> which sometimes results in an ape-like appearance.

Maxillomandibular advancement: Consider for patients with hypopharyngeal narrowing, veloorohypopharyngeal narrowing, and retrognathia; patients with BMI < 30 kg/m² may have better surgical outcomes.

### What surgical OSA treatment targets neuromuscular tone?

Hypoglossal nerve stimulation This pacemaker-like device is connected to a wire that attaches a small cuff to the hypoglossal nerve. 43,44 When turned on, the device stimulates the hypoglossal nerve and protrudes the tongue, opening the pharyngeal airway and maintaining patency. 43,44 Hypoglossal nerve stimulation reduces the AHI by 68%, from 29.3 events to 9 events per hour, 43 with sustained benefit at 3-year follow-up. 44 Participants with a BMI of >32 kg/m² and those with concentric retropalatal collapse on drug-induced sleep endoscopy are not candidates for hypoglossal nerve stimulation. 43

Hypoglossal nerve stimulation: Consider for patients with BMI < 32 kg/m<sup>2</sup>; drug-induced sleep endoscopy is required to determine the patient's candidacy; concentric retropalatal collapse is a contraindication.

# What treatments exist for OSA patients who do not want any device or surgery?

Weight loss Patients with OSA and comorbid obesity should be counseled on long-term weight management. A goal BMI <25 kg/m² through dietary or surgical weight loss may improve the AHI in obese patients with OSA.<sup>45</sup> Major weight loss in conjunction with PAP therapy may allow for reduced PAP requirements,<sup>45</sup> which may facilitate patient comfort. Weight loss may also benefit management of other existing comorbidities such as hypertension and high cholesterol.<sup>45</sup>

Weight loss: Consider for all overweight or obese patients with OSA.

**Positional therapy** Positional therapy was developed to keep patients in a nonsupine position. Supine position can decrease the area of the upper airway and worsen severity of OSA. Avoidance of supine sleep, with isolated lateral positioning, could potentially increase airway patency. <sup>46</sup> Patients who pursue positional therapy are advised to wear a device, either a commercially available product or homemade positioner, to prevent supine sleep.

Positional therapy: Consider avoidance of supine sleep for patients whose OSA is worse while back-sleeping.

*Nasal expiratory PAP* Nasal expiratory PAP is a Food and Drug Administration—cleared device that relies on patent nasal passages. A randomized controlled trial showed a 42.7% decrease in AHI with use of the device.<sup>47</sup> The device valve rests in the nose and acts as a one-way resistor, permitting unobstructed inspiration.<sup>48</sup> Airflow during expiration increases resistance, which creates expiratory PAP during expiration that is maintained until the start of the next inspiration.<sup>48</sup>

Nasal expiratory PAP: May consider for patients with patent nasal passages who do not mouth-breathe during sleep; less studied compared to other OSA treatments.

**Noninvasive oral pressure therapy** Oral pressure therapy is delivered via a device that rests within the mouth. The device produces a negative pressure with a vacuum within the oral cavity, which draws the soft palate forward and stabilizes the airway. Although one study suggested significant improvement in AHI with use of oral pressure therapy, this benefit was not seen among all groups of patients in the study. 49

Noninvasive oral pressure therapy: May consider for patients who can tolerate the device; less studied compared to other OSA treatments.

### CONCLUSION

Patients with neurologic disease who report fatigue and tiredness may have comorbid OSA. Diagnosis and treatment of OSA may contribute to improved quality of life for patients with neurologic disease. Continuous positive airway therapy continues to be the gold standard OSA treatment, and other modalities of PAP delivery are also available. Patients who cannot use or do not tolerate CPAP may require a multidisciplinary approach to care with collaboration among neurologists, sleep medicine specialists, dentists, and surgeons to determine the most appropriate

management. This tailored approach can help identify an individual patient's specific sites of airway obstruction and develop a treatment plan that may maximize benefit to the patient.

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### Take-home points

- Positive airway pressure (PAP) is the firstline treatment for obstructive sleep apnea (OSA). Multiple varieties of PAP delivery are available for clinical use.
- Mandibular advancement devices may be considered for appropriate patients with mild to moderate OSA or patients with severe OSA who decline or cannot use PAP therapy.
- Nasal, palatal, or jaw surgeries may be considered for treatment of OSA in appropriate patients.
- Hypoglossal nerve stimulation targets the impaired neuromuscular tone implicated in OSA.
- Weight loss should be considered in all appropriate OSA patients as part of a multimodal treatment approach.

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