




Evaluation and Management of Children with Obstructive Sleep Apnea Syndrome

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Abstract

Obstructive sleep apnea syndrome (OSAS) is a common pediatric disorder characterized by recurrent events of partial or complete upper airway obstruction during sleep which result in abnormal ventilation and sleep pattern. OSAS in children is associated with neurobehavioral deficits and cardiovascular morbidity which highlights the need for prompt recognition, diagnosis, and treatment. The purpose of this state-of-the-art review is to provide an update on the evaluation and management of children with OSAS with emphasis on children with complex medical comorbidities and those with residual OSAS following first-line treatment. Proposed treatment strategies reflecting recommendations from a variety of professional societies are presented. All children should be screened for OSAS and those with typical symptoms (e.g., snoring, restless sleep, and daytime hyperactivity) or risk factors (e.g., neurologic, genetic, and craniofacial disorders) should undergo further evaluation including referral to a sleep specialist or pediatric otolaryngologist and overnight polysomnography, which provides a definitive diagnosis. A cardiology and/or endocrinology evaluation should be considered in high-risk children. For the majority of children, first-line treatment is tonsillectomy with or without adenoidectomy; however, some children exhibit multiple levels of airway obstruction and may require additional evaluation and management. Anti-inflammatory medications, weight loss, and oral appliances may be appropriate in select cases, particularly for mild OSAS. Following initial treatment, all children should be monitored for residual symptoms and polysomnography may be repeated to identify persistent disease, which can be managed with positive airway pressure ventilation and additional surgical approaches if required.

Keywords Adenotonsillectomy (AT) · Diagnosis · Obstructive sleep apnea syndrome (OSAS) · Pediatrics · Polysomnography · Sleep disordered breathing (SDB)

Introduction

Sleep disordered breathing (SDB) in children refers to a spectrum of disorders which occur during sleep including central apnea, hypoventilation, and obstructive hypoventilation. The most severe manifestation of obstructive hypoventilation, obstructive sleep apnea syndrome (OSAS), is characterized by recurrent events of partial or complete upper airway obstruction during sleep which leads to abnormal ventilation and sleep pattern [1, 2]. The prevalence in

children is 2–4% [3] and is increasing with the rising trend of childhood obesity [4]. OSAS is associated with neurobehavioral deficits [5], cardiovascular morbidity [6], poor quality of life [7], and increased health care utilization [8] which underscores the importance of detection and treatment. Consequently, the American Academy of Pediatrics (AAP) recommends screening for OSAS at routine medical visits. The diagnosis should be considered in children with typical symptoms (e.g., snoring, restless sleep, or daytime hyperactivity) or risk factors (e.g., craniofacial, neurologic, or genetic disorders) [9] and is confirmed with overnight polysomnography.

While the pathophysiology of pediatric OSAS is multifactorial, the most common cause is overgrowth of the adenoid and tonsils leading to upper airway restriction during sleep. Accordingly, first-line treatment is tonsillectomy with or without adenoidectomy, collectively referred to as adenotonsillectomy (AT) [1, 10]. Respiratory parameters improve

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following AT in the majority of children, but 40–75% have some level of persistent disease [11–13]. When AT is not indicated, if there is residual OSAS following surgery, or when medical comorbidities are present, additional evaluation and a modified management plan is often required. This may involve positive airway pressure (PAP) ventilation, weight management, dental procedures, secondary airway surgery, or medical therapy. Choice of treatment is tailored to the individual child depending on their comorbidities, preferences, and disease severity and is planned in consultation with a specialist in pediatric otolaryngology or sleep medicine.

Evaluation

A proposed strategy for the evaluation of children with suspected OSAS is presented in Table 1.

History and Physical Examination

Clinical History

The most common complaints reported by parents of children with OSAS are snoring and difficulty breathing during sleep. Although almost all children with OSAS snore, snoring has low specificity for OSAS and clinical symptoms

Table 1 Evaluation of the child with suspected OSAS or residual OSAS following AT

Otherwise healthy children
Clinical evaluation: history and physical exam (recommended by AAP [1], AASM [114], AAO-HNS [10], and ERS [115])
Diagnostic polysomnogram (recommended by AAP [1], AASM [26, 114], and ERS [115])
<ul style="list-style-type: none"> • AAO-HNS recommends polysomnography if the need for surgery is unclear or if there is discordance between the clinical history and the tonsillar size on physical exam [28]
Alternative testing when polysomnography is not available
<ul style="list-style-type: none"> • AAP: options include nocturnal video recording, nocturnal oximetry, daytime nap polysomnography, or ambulatory polysomnography [1] • ERS: options include ambulatory polysomnography or polygraphy, nocturnal oximetry, Pediatric Sleep Questionnaire, or Sleep Clinical Record [115] • AASM: does not recommend home sleep apnea testing for OSAS diagnosis in children [37] • AAO-HNS: home-based studies may be considered but are not recommended for routine use [28]
Consider endocrinology/weight management referral for obese children
Consider cardiology referral for children with severe sleep apnea or cardiometabolic risk factors
Children with complex comorbidities
<i>Includes children with obesity, neuromuscular disorders, craniofacial abnormalities, genetic syndromes, sickle cell disease, etc</i>
Clinical evaluation: history & physical exam (recommended by AAP [1], AASM [114], AAO-HNS [10], and ERS [115])
Diagnostic polysomnogram (recommended by AAP [1], AASM [114], AAO-HNS [10], and ERS [115])
Expert evaluation (e.g., pediatric pulmonology, pediatric otolaryngology, sleep medicine, craniofacial team)
Consider endocrinology/weight management referral for obese children
Consider cardiology referral for children with severe sleep apnea or cardiometabolic risk factors
Children with residual OSAS following AT
Posttreatment reevaluation of high-risk patients
<ul style="list-style-type: none"> • AAP: High-risk children (defined as those with significantly abnormal baseline polysomnogram, sequelae of OSAS, obesity, or continued symptoms following treatment) should be reevaluated with objective testing or referred to a sleep specialist [1] • AASM: Polysomnography should be repeated following adenotonsillectomy in children at risk of recurrence: mild pre-operative OSAS with residual symptoms following surgery, pre-operative evidence of moderate–severe OSAS, obesity, craniofacial abnormalities, or neurologic disorders [114] • ERS: Patients at risk of persistent OSAS or with persistent symptoms should be reevaluated with polysomnography or polygraphy [115]
Expert evaluation (pediatric pulmonology, sleep medicine, pediatric otolaryngology, endocrinology)
Polysomnography titration for positive airway pressure therapy (AAP [1])
Nasopharyngoscopy or drug-induced sleep endoscopy
Imaging (computed tomography, dynamic magnetic resonance imaging)
Cardiology referral for persistent, untreated OSAS

AAO-HNS American Academy of Otolaryngology—Head & Neck Surgery, AAP American Academy of Pediatrics, AASM American Academy of Sleep Medicine, ERS European Respiratory Society, OSAS obstructive sleep apnea syndrome

alone cannot reliably distinguish OSAS from primary snoring [14, 15]. Other symptoms during sleep include paradoxical breathing (in which the thoracic wall moves inward with respiration), gasping, restless sleep, frequent awakenings, or respiratory pauses. The duration and frequency of these symptoms should be elicited as well as the degree of persistence (e.g., intermittent, only during illness, or all the time). Other common nocturnal findings include diaphoresis and enuresis. Daytime manifestations of OSAS in children differ from adults. Excessive daytime sleepiness is uncommon [16, 17]. Neurobehavioral sequelae are more often present and are thought to be the result of chronic exposure to intermittent hypoxemia and sleep deprivation. These impairments fall into two main categories: behavioral (such as aggression, impulsivity, and hyperactivity) [18] and neurocognitive (with associated deficits in attention, executive function, and language) [19]. Cognitive deficits correlate with OSAS severity in a dose-dependent manner and are manifest even with mild OSAS [20]. A comprehensive evaluation of sleep hygiene should be performed including a discussion of the frequency of napping, timing of sleep, sleep environment, large evening meals, and late-night screen use.

OSAS is more prevalent in children with certain comorbid conditions including but not limited to those displayed in Table 2.

Physical Examination

The physical examination begins with a general observation of the child which may reveal mouth breathing, adenoidal facies, or dysmorphism suggestive of a genetic syndrome. Voice quality may be hyponasal due to nasal obstruction or muffled secondary to tonsillar enlargement. Tonsil size is graded using the Brodsky score [21]. The head and neck exam may reveal other clues to the source of airway obstruction such as retrognathia, micrognathia, midfacial hypoplasia, nasal obstruction, and macroglossia. Although cardiac abnormalities are uncommon and usually manifest in the context of longstanding severe disease, children can develop systemic hypertension or pulmonary hypertension [22–24]. While difficult to detect clinically, the latter may manifest with pronounced closure of the pulmonary valve (loud P2). In young infants and in children with a non-contributory physical examination, neurological impairments that affect upper airway muscle tone should be excluded. The child's growth should also be evaluated, as obesity is a risk factor.

Polysomnography

The gold standard to diagnose and assess the severity of OSAS in children is overnight polysomnography. Guidelines for performing polysomnography are published by the

Table 2 Medical conditions associated with OSAS

Craniofacial anomalies
Achondroplasia
Apert syndrome
Beckwith-Wiedemann
Choanal stenosis
Crouzon syndrome
Down syndrome
Goldenhar syndrome
Hallerman-Streiff syndrome
Klippel-Feil syndrome
Laryngomalacia
Marfan syndrome
Micrognathia
Midface hypoplasia
Nager syndrome
Palate formation (e.g., high and narrow hard palate)
Pfeiffer syndrome
Pierre Robin sequence
Retrognathia
Rubinstein-Taybi
Stickler syndrome
Subglottic stenosis
Treacher-Collins syndrome
Neurological disorders
Arnold-Chiari malformation
Cerebral palsy
Möbius syndrome
Muscular dystrophy
Myasthenia gravis
Myelomeningocele
Myotonic dystrophy
Poliomyelitis
Spinal muscular atrophy
Syringobulbia
Syringomyelia
Miscellaneous disorders
Airway papillomatosis
Congenital hypothyroidism
Face and neck burns
Family history of obstructive sleep apnea syndrome
History of prematurity
Mucopolysaccharidoses (e.g., Hurler, Hunter)
Obesity
Polycystic ovary syndrome
Prader Willi syndrome
Sickle cell disease
Postoperative disorders
Postadenotonsillectomy leading to naso- and/or oro-pharyngeal stenosis
Postpharyngeal flap: leading to naso- and/or oro-pharyngeal stenosis

American Academy of Sleep Medicine (AASM) [25]. The parameters monitored include but are not limited to electroencephalogram derivations, electrocardiogram tracing, oxygen saturation, end-tidal CO₂ (P_{ETCO₂}), body position and movement, sleep staging and architecture, and apneas (obstructive, mixed, and central).

The criteria for pediatric OSAS are shown in Table 3 and published in the *International Classification of Sleep Disorders, 3rd Ed.* [26]. The diagnosis requires both (A) signs or symptoms of SDB and (B) abnormal polysomnogram findings. Clinical findings which fulfill the first criterion include snoring; labored, paradoxical, or obstructed breathing during sleep; and daytime consequences (sleepiness/hyperactivity). The polysomnographic criterion for diagnosis requires either (1) one or more obstructive events (obstructive or mixed apnea or obstructive hypopnea) per hour of sleep or (2) obstructive hypoventilation, manifested by P_{ETCO₂} > 50 mm Hg for > 25% of sleep time and/or arterial oxygen desaturation coupled with snoring, paradoxical

thoracoabdominal movement, or flattening of the nasal airway pressure waveform [26]. Patients up to age 18 may be scored using the diagnostic criteria for children, but sleep specialists can choose to score children ≥ 13 years using adult criteria [25]. Particularly in very young children, central apnea may co-exist with OSAS. Careful scoring of the polysomnogram is needed to distinguish between central and obstructive events as the etiology and management of central apnea differs from OSAS. Several polysomnographic respiratory parameters of importance and their definitions are presented in Table 4. For further detail the reader is directed to the *AASM Manual for the Scoring of Sleep and Associated Events* [25].

There is no universally accepted classification for OSAS severity in children, although many studies use the apnea hypopnea index (AHI) to categorize OSAS as mild (AHI 1–4.9), moderate (AHI 5–9.9), or severe (AHI > 10) [10]. A cutoff of AHI > 30 to denote very severe OSAS has also been proposed [27]. Other objective findings of disease severity

Table 3 Diagnostic Criteria for Pediatric OSAS Based on AASM *International Classification of Sleep Disorders*, 3rd ed [26]. (Diagnosis Requires A + B)

A. The presence of one or more of the following:

1. Snoring
2. Labored, paradoxical, or obstructed breathing during the child's sleep
3. Sleepiness, hyperactivity, behavioral problems, or learning problems

B. Polysomnography demonstrates one or more of the following:

1. One or more obstructive or mixed apneas, or obstructive hypopneas, per hour of sleep
2. A pattern of obstructive hypoventilation, defined as at least 25% of total sleep time with hypercapnia (*P_{ETCO₂} > 50 mm Hg) and/or arterial oxygen desaturation in association with one or more of the following: snoring, flattening of the inspiratory nasal pressure waveform, and paradoxical thoracoabdominal motion

* P_{ETCO₂}: end-tidal CO₂

Table 4 Pediatric respiratory events derived from polysomnography. Modified from The AASM Manual for the Scoring of Sleep and Associated Events [25]

Obstructive apnea: At least a 90% drop in peak signal excursion of the oronasal thermal sensor in the presence of continued respiratory effort lasting longer than 2 respiratory cycle times

Central apnea: At least a 90% drop in peak signal excursion of the oronasal thermal sensor in the absence of respiratory effort and at least one of the following: (1) ≥ the event lasts 20 s; (2) the event lasts at least 2 respiratory cycles and is associated with ≥ 3% oxygen desaturation; or (3) the central event lasts at least 2 respiratory cycle times and is associated with heart deceleration

Mixed apnea: Criteria for apnea need to be met for at least 2 respiratory cycles and associated with absent respiratory effort during one portion of the event and inspiratory effort in another portion

Hypopnea: All the following three criteria need to be met: (1) at least 30% decrease in the amplitude of the nasal pressure signal; (2) duration ≥ 2 respiratory cycles times; and (3) ≥ 3% oxygen desaturation or the event is associated with arousal

Obstructive hypopnea: To score hypopnea as obstructive, the following need to be met: (1) snoring during the event; (2) increased inspiratory flattening of the nasal pressure flow signal compared with baseline breathing; and (3) associated thoracoabdominal paradox occurs during the event but not during pre-event breathing

Hypoventilation: Criteria are met when end-tidal CO₂ (P_{ETCO₂}) > 50 mmHg persists for more than 25% of total sleep time

Apnea index (AI): Number of obstructive and/or central apneic events per hour of sleep

Obstructive apnea index (oAI): Number of obstructive apneic events per hour of sleep

Hypopnea index (HI): Number of hypopneas per hour of sleep

Apnea/hypopnea index (AHI): The summation of apnea index and hypopnea index

Obstructive apnea/hypopnea index (oAHI): The summation of obstructive apneic events and hypopneic events per hour of sleep

include presence and length of oxygen desaturations, degree of hypoventilation, sleep fragmentation, and decreased total sleep time. It is important to note that the severity of polysomnogram findings does not always correlate with the degree of morbidity and thus treatment decisions should take into account the child's symptoms and relevant comorbidities in addition to the objective polysomnogram findings.

It should be mentioned that there is controversy regarding indications for polysomnography in children. Guidelines published by the AASM, AAP, and European Respiratory Society (ERS) recommend polysomnography for all children when the clinical assessment is suggestive of OSAS (Table 1). In contrast, the American Academy of Otolaryngology—Head & Neck Surgery favors a more limited use of polysomnography and recommends routine use only in high-risk populations (children with age < 2, obesity, Down syndrome, craniofacial abnormalities, neuromuscular disorders, sickle cell disease, or mucopolysaccharidoses) or if the need for surgery is uncertain or there is discordance between the history of SDB and the tonsillar size on physical exam [28].

Screening Tools

Due to the scarcity of accredited sleep laboratories and significant resources required to perform polysomnography, there has been considerable effort invested into the development of alternative screening or diagnostic methods which could be more widely available.

Questionnaires

Owing to their convenience, a reliable screening questionnaire to identify pediatric OSAS would have significant clinical utility. A number of clinical scoring instruments have been developed including the OSA score [29], pediatric sleep questionnaire (PSQ) [30], and OSA-18 [31], among others [32]. The diagnostic accuracy varies depending on the cutoff score and study population but is generally too low to be considered an alternative diagnostic method. While they cannot accurately identify OSAS, questionnaires do have clinical utility in assessing the impact of OSAS on quality of life. For example, the PSQ showed stronger correlation to behavior, sleepiness, and quality of life metrics than polysomnogram parameters, leading the authors to suggest that PSQ score is a useful adjunct to polysomnography [33]. Other groups have developed predictive models using a combination of symptoms-based questions and physical exam findings or results from home sleep apnea testing (HSAT) [34, 35], but their utility in clinical practice has not been determined.

Home Sleep Apnea Testing and Pulse Oximetry

HSAT is performed in the patient's home and monitors fewer physiologic variables than full polysomnography. In theory, HSAT could have significant benefits over laboratory-based polysomnography in terms of cost, resources, and disruption to family life. HSAT is an acceptable method for the diagnosis of OSAS in uncomplicated adult patients [36]; however, an AASM task force found insufficient evidence to support the use of HSAT in children [37]. Further validation studies are required to assess the feasibility and validity of these studies and determine their optimal role in pediatric OSAS.

Continuous pulse oximetry has also been studied as a screening or diagnostic tool when polysomnography is not available. Although intermittent oxygen desaturations on overnight continuous pulse oximetry is highly suggestive of OSAS [38], not all children with OSAS exhibit nocturnal hypoxemia [39] and, therefore, children with negative studies require follow-up polysomnography to rule out OSAS. A large prospective study recently evaluated the diagnostic utility of nocturnal oximetry to estimate the AHI of habitually snoring children [40]. An automatic analysis algorithm achieved an accuracy of 75.2, 81.7, and 90.2% in diagnosing children with AHI ≥ 1 , ≥ 5 , and ≥ 10 , respectively, raising the possibility that overnight oximetry could play an important role in the diagnosis of pediatric OSAS especially in places where polysomnography is not available [40].

Upper Airway Evaluation

Radiographic Evaluation

Several radiological techniques are available to evaluate the upper airway including lateral neck radiographs, cephalometrics, fluoroscopy, computerized tomography (CT), and magnetic resonance imaging (MRI) [41–44]. These modalities are most often used in the research setting to study the pathophysiology of OSAS, though a small number of studies have explored their diagnostic potential with varying results [45]. In clinical practice, radiographic techniques are typically reserved for upper airway evaluation in children with comorbid conditions such as craniofacial anomalies or neurological conditions and may assist in identification of level(s) of obstruction and treatment selection. Conventional techniques for upper airway evaluation such as fluoroscopy, lateral neck cephalometry, and CT have substantial limitations as they evaluate neck anatomy in the awake state (and thus may not demonstrate dynamic airway collapse occurring in sleep) and expose children to radiation. In light of these limitations, dynamic MRI and drug-induced sleep endoscopy (DISE) have emerged as alternative methods of assessment.

Dynamic MRI

MRI is particularly powerful since it may be used to reconstruct the upper airway (including soft tissues and skeletal structure) [43, 46] in three dimensions and to evaluate upper airway dynamics [47, 48]. Dynamic MRI captures dynamic changes of the upper airway throughout the respiratory cycle and is typically used for children with persistent OSAS following AT or for high-risk children with multiple sites of upper airway obstruction [49]. Identified sites of obstruction may then be targeted surgically. Reports regarding MRI-directed surgical outcomes are sparse; however, a meta-analysis including 68 patients found that the most common procedure performed after dynamic MRI was lingual tonsillectomy (57%) and the mean AHI reduction was 7.4 events/h [50]. At present dynamic MRI is not widely used and is only performed at a small number of specialized sleep centers.

Drug-Induced Sleep Endoscopy (DISE)

In DISE, the upper airway is evaluated with a flexible endoscope while the child is in a pharmacologically induced sleep-like state [51]. This method enables dynamic evaluation of the airway and identification of areas of obstruction which may become surgical targets. DISE-directed procedures include lingual tonsillectomy, supraglottoplasty, nasal surgery, revision adenoidectomy, uvulopalatopharyngoplasty,

and partial midline glossectomy [52]. Despite its increasingly widespread use, studies evaluating outcomes after DISE-directed therapy are sparse. A small retrospective study found that approximately half of children undergoing DISE-directed surgery experienced resolution of OSAS; however, in the subgroup of children who underwent DISE for residual OSAS following AT, there was no improvement in obstructive AHI [52]. A small prospective study of 20 children reported a decrease in mean AHI from 6.1 to 1.9 following DISE-directed surgery [53]. Although use of DISE continues to grow, larger and more rigorous studies should be performed in order to evaluate outcomes associated with DISE-directed surgical management and identify subpopulations who benefit most from this approach.

Management

A proposed strategy for the management of children with OSAS is presented in Table 5.

Expert Evaluation

The link between OSAS and cardiometabolic morbidity in adults is well established [54], and current evidence also

Table 5 Management of the Child with OSAS

Otherwise Healthy Children		
Mild OSAS	Moderate OSAS	Severe OSAS
<ul style="list-style-type: none"> • AT vs watchful waiting for 6 months • Intranasal corticosteroid • Weight management, if overweight/obese 	<ul style="list-style-type: none"> • Specialist referral (e.g., sleep medicine, otolaryngology) • AT • PAP therapy • Weight management, if overweight/obese 	<ul style="list-style-type: none"> • Specialist referral (e.g., sleep medicine, otolaryngology) • Cardiology evaluation • AT with overnight observation • PAP therapy • Weight management, if overweight/obese
Children with complex comorbidities		
Mild OSAS	Moderate OSAS	Severe OSAS
<ul style="list-style-type: none"> • Specialist referral (e.g., sleep medicine, otolaryngology) • AT • Intranasal corticosteroid • Weight management, if overweight/obese 	<ul style="list-style-type: none"> • Specialist referral (e.g., sleep medicine, otolaryngology) • AT with overnight observation • Other airway and/or craniofacial surgery • PAP therapy • Weight management, if overweight/obese 	<ul style="list-style-type: none"> • Specialist referral (e.g., sleep medicine, otolaryngology, craniofacial team) • Cardiology evaluation • AT with overnight observation • Other airway and/or craniofacial surgery • PAP therapy • Weight management, if overweight/obese
Children with residual OSAS following AT		
Mild OSAS	Moderate OSAS	Severe OSAS
<ul style="list-style-type: none"> • Intranasal corticosteroid • Weight management, if overweight/obese 	<ul style="list-style-type: none"> • PAP therapy • Weight management, if overweight/obese 	<ul style="list-style-type: none"> • PAP therapy • Weight management, if overweight/obese • Secondary surgical intervention • Tracheostomy

AT adenotonsillectomy, OSAS obstructive sleep apnea syndrome, PAP positive airway pressure

supports this relationship in the pediatric population [6]. Accordingly, children with severe OSAS and those with pre-existing risk factors for cardiometabolic disease may require evaluation by a specialist. Children with severe OSAS are often referred to cardiology prior to surgery, as are children with overt cardiac manifestations (such as elevated blood pressure or signs of pulmonary hypertension). Children who are obese may also benefit from an endocrinology evaluation.

Surgical Management

Adenotonsillectomy

In healthy children over two years of age with adenotonsillar hypertrophy, first-line treatment of OSAS is AT [1, 10]. This recommendation is supported by the results of the Childhood Adenotonsillectomy Trial (CHAT), a large randomized controlled trial which found that surgical intervention with AT was associated with significant improvements in behavior, quality of life, and polysomnographic findings as compared to watchful waiting. At seven months follow-up, polysomnogram findings normalized in 79% of surgical patients compared to 49% in the watchful waiting arm [7]. In a meta-analysis of three studies (including CHAT), Chinnadurai et al. reported a 4.8-point improvement in AHI for patients undergoing surgical intervention compared to watchful waiting [55].

Classically, tonsillectomy was performed using “cold” techniques to dissect the tonsillar capsule off the pharyngeal constrictor muscles. Alternatives to conventional cold dissection include electrosurgery (mono- or bi-polar) and plasma-mediated ablation (coblation) [56]. Partial tonsillectomy, a modified procedure in which the majority of tonsillar tissue is removed but the tonsillar capsule remains in place, is increasingly being utilized to treat OSAS in children [57]. Complication rates of partial tonsillectomy (also known as tonsillotomy and intracapsular tonsillectomy) are lower compared to standard tonsillectomy; however, few studies have examined its therapeutic effect [57]. The subset of children with both obesity and asthma are more likely to experience resolution of OSAS following traditional tonsillectomy compared to partial tonsillectomy [58], but these findings should be evaluated in prospective trials. Immediately following tonsillectomy, children should be monitored for respiratory complications which range in severity from mild oxygen desaturation to postoperative pulmonary edema. Other complications include throat pain, bleeding, nausea/vomiting, dehydration, delayed feeding, velopharyngeal insufficiency, and rarely death [10]. Most tonsillectomies are performed as outpatient procedures although children with age < 3 or severe OSAS (AHI > 10 or oxygen saturation

nadir < 80%) are at increased risk for complications and should be observed overnight [10].

AT is generally successful in improving respiratory parameters but is not curative in all cases. In a multicenter study of almost 600 children undergoing AT for OSAS, AHI decreased from 18.2 to 4.1, but complete resolution of OSAS was observed in only 27.2% of children [12]. Risk factors for persistence include age > 7, severe disease, chronic asthma, and obesity [12, 59].

Other Upper Airway Surgeries

Surgical procedures beyond AT are typically reserved for children with comorbidities leading to airway obstruction at multiple levels or for children with persistent OSAS following AT. Potential causes of airway obstruction include turbinate hypertrophy, nasal septal deviation, redundant palatal tissue, large tongue base, retrognathia, regrowth of the adenoids, laryngomalacia, or lingual tonsil hypertrophy [60]. Obstruction at the level of the tongue base can be addressed using a number of procedures including lingual tonsillectomy and posterior midline glossectomy. Obstruction may also be addressed at the level of the palate (uvulopalatopharyngoplasty [61]), lateral pharyngeal walls (expansion sphincter pharyngoplasty [62]), nasal cavity (turbinate reduction [63]), or adenoids (revision adenoidectomy [64]).

When airway obstruction is severe due to craniofacial disorders restricting the upper airway, more extensive surgical approaches are used. In children with mandibular insufficiency as seen in Pierre-Robin Sequence, Treacher-Collins Syndrome, and other causes of congenital retrognathia, mandibular advancement or mandibular distraction osteogenesis can effectively alleviate upper airway obstruction and avoid tracheostomy dependence [65, 66]. Genioglossus advancement and hyoid suspension have also been used to treat OSAS, though the body of evidence is small and most studies involve adults [67, 68].

Tracheotomy

Tracheotomy is the definitive surgical treatment for OSAS. The majority of patients who require tracheotomy have either severe craniofacial abnormalities or neuromuscular conditions causing hypotonia [69]. Although many children remain tracheostomy-dependent, the procedure can also be used as a temporary measure to manage severe OSAS while awaiting other surgical treatment.

Non-surgical Management

Positive Airway Pressure (PAP)

Children with small tonsils/adenoids and those who are not surgical candidates or whose parents decline surgery should be managed with PAP therapy during sleep. PAP therapy maintains airway patency by serving as a pneumatic splint for the soft tissues of the upper airway leading to increased cross-sectional airway area throughout the respiratory cycle. Initiation of PAP therapy begins with an overnight sleep study with PAP titration, ideally in an accredited sleep laboratory or in a hospital where an expert technician or respiratory therapist can monitor respiratory and sleep parameters. Pressure adjustments are performed with the goal of eliminating respiratory events and correcting gas exchange abnormalities, while also maintaining patient tolerability. Technical details regarding PAP titration may be found elsewhere [70]. Continuous PAP (CPAP) and bilevel PAP (BPAP) are equally effective, although CPAP is used most frequently. PAP therapy in children improves both respiratory parameters and neurobehavioral sequelae of OSAS such as attention span, hyperactivity, daytime sleepiness, and school performance [71, 72]. Studies in adults have demonstrated improvement in blood pressure and cognitive function [73, 74], but data regarding the corollary outcomes in children are limited.

Although PAP therapy is an effective treatment modality for children of all ages and with a range of underlying pathophysiological mechanisms, effectiveness is limited by poor adherence in some studies. In a prospective study on PAP compliance in children, Marcus and colleagues noted a high dropout rate (35%), low average nightly use (5.3 h/night), and frequent overreporting of adherence by parents as compared to objective compliance data [71]. Discomfort associated with the face mask is frequently reported and likely contributes to poor adherence. Compliance can be significantly improved by close clinical follow-up in addition to intensive education [75] and gradual introduction of the PAP mask [76]. Side effects include nasal congestion, oronasal dryness, epistaxis, eye irritation from air leak, facial pain, and skin abrasion [77]. Though rare, skin necrosis and ulceration have been reported [78]. Long-term CPAP use has also been linked to midface hypoplasia [79] and midfacial retrusion over time [80]. These skeletal changes appear to be common, as global facial flattening and maxillary retrusion can be observed in 68% and 37% of patients, respectively [81]. Accordingly, routine follow-up with special attention to midface growth is essential in children when long-term PAP therapy is anticipated.

Medical Therapy

Medical therapies have been proposed as an alternative to surgery, especially in children with mild OSAS. A six-week course of once-daily intranasal budesonide has been shown to improve quality of life and respiratory parameters measured on polysomnography with a sustained effect two months after discontinuation [82, 83], and a recent meta-analysis found that montelukast improved the AHI by 55% when used alone and by 70% when used in conjunction with intranasal corticosteroids in children with mild OSAS [84]. These therapies hold promise as potential treatments for mild uncomplicated OSAS in children. Of note, obese children and those with age > 7 are less likely to benefit from anti-inflammatory therapy [85].

Dental Procedures

Rapid maxillary expansion is an orthodontic procedure which expands the airway by increasing the width of the hard palate using a dental device secured over the maxillary teeth and an accompanying expansion screw. Two studies have shown normalization of polysomnogram parameters and symptoms using this technique in children with maxillary restriction and dental malocclusion [86, 87]. Oral appliances including tongue devices and mandibular advancement devices move the tongue and mandible forward and away from posterior pharynx to improve upper airway patency. In a small group of children with mild OSAS and dysgnathia, oral jaw-positioning appliances were shown to reduce AHI, improve sleep, and alleviate daytime sleepiness [88]. A recent Cochrane review of randomized studies for oral appliances in pediatric OSAS identified only this one study and concluded there is insufficient evidence to support or refute their utility [89].

Special Considerations

Certain children with OSAS present unique challenges to treatment and their management warrants further discussion.

Residual OSAS After Adenotonsillectomy (AT)

Not all children experience resolution of OSAS following AT and some (including those with chromosomal, craniofacial, or neuromuscular disorders) are at increased risk of persistence [12, 59, 90]. Residual OSAS may be identified by repeat polysomnogram (which is recommended for those with risk factors) or by recurrence of clinical symptoms. In the latter case, the diagnosis should be confirmed with polysomnography. Options for the management of persistent

OSAS include airway evaluation with possible surgical intervention, PAP, and medical therapy. DISE or dynamic MRI can be used to evaluate the upper airway and identify sites of persistent obstruction which can be targeted surgically, as discussed above. PAP is effective in this setting, although difficulties with tolerability and poor adherence still apply [72]. A combination of montelukast and intranasal budesonide was also shown to improve AHI and oxygen nadir in children with mild OSAS following AT [83]. Lastly, weight management should be recommended for children who are overweight or obese (see next section).

Children with Complex Comorbidities

Obesity

Obesity is reaching epidemic proportions across all age groups, including children, and increases the risk for OSAS with an odds ratio of 4.5 [91]. Some studies have reported the prevalence of OSAS to be as high as 45% in obese children [92]. The cause of this association is not fully elucidated, but adenotonsillar hypertrophy, increased airway collapsibility, alterations in body fat composition, and abnormal ventilatory response likely each play a role [4]. If adenotonsillar hypertrophy is present, AT is first-line treatment and significantly improves OSAS symptoms and polysomnogram parameters; however, persistent OSAS is identified in 33–76% of obese children compared to 15–37% of non-obese children [93]. While the reason for high failure rates is unclear, residual adenoid tissue and increased soft palate volume have been observed in obese children following AT and may contribute to persistent airway obstruction [94]. PAP is a good treatment alternative for children without adenotonsillar hypertrophy or those with residual OSAS following AT. A small number of studies have explored the hypothesis that weight loss may improve OSAS in obese children. Andersen and colleagues prospectively enrolled 62 overweight or obese children with AHI > 2 in an individualized, family-centered obesity treatment protocol. After one year, the AHI normalized in 44% of children and the change in BMI standard deviation score was associated with the decrease in AHI; however, most of the children (71%) had mild OSAS at baseline [95]. Although the effect of weight loss on OSAS outcomes is not well studied, weight management is routinely recommended. A decrease in BMI is associated with improved metabolic outcomes in obese children and should be routinely encouraged notwithstanding its potential benefits regarding OSAS [96]. Lastly, further surgical management may be considered although studies reporting outcomes of secondary surgery such as uvulopalatopharyngoplasty or supraglottoplasty in this population are sparse and have so far demonstrated poor efficacy [97].

Down Syndrome

The prevalence of OSAS among children with Down syndrome may be as high as 70% and almost half have severe disease [98]. Accordingly, the AAP recommends routine screening for SDB/OSAS in children with Down syndrome [99]. The pathophysiology of OSAS in this population is multifactorial and includes hypoplasia of the midface/mandible, relative macroglossia, lingual tonsil hypertrophy, comorbid laryngomalacia, obesity, decreased airway tone, and hypopharyngeal collapse [100–102]. Recognition and treatment of OSAS is particularly important as the consequences of untreated OSAS may exacerbate underlying cardiac disease, behavioral problems, and cognitive challenges [103–105]. The pathophysiology, morbidity, and treatment of OSAS in patients with Down syndrome was reviewed recently and highlights the need for further research regarding treatment options and outcomes in this high-risk population [106]. AT is first-line treatment but has a low success rate of approximately 10–20% [24, 107]. Residual OSAS may be treated with PAP or further surgery. DISE-directed therapy resulted in improvements in polysomnogram parameters in a small cohort of children with Down syndrome, but the study was not powered to detect statistically significant differences [108].

Craniofacial Disorders

Children with disorders associated with midfacial hypoplasia and/or mandibular deficiency deserve special attention in their evaluation and management. We discuss here Pierre-Robin sequence, however, a similar approach may be applied to similar disorders (see Tables 2, 5). Pierre-Robin sequence is characterized by a triad of micrognathia, glossoptosis, and upper airway obstruction with or without cleft palate and can occur as an isolated finding or in the context of other congenital abnormalities or syndromes. The presence of micrognathia with glossoptosis significantly increases the risk of developing OSAS and studies have reported a high prevalence (46–83%) of OSAS [109]. The severity of upper airway obstruction and presentation of SDB varies widely; some children experience respiratory distress requiring intubation at birth, while others present later in life with poor weight gain [110]. Initial management with lateral or prone positioning and placement of a nasopharyngeal airway relieves the obstruction in some children [111]. A trial of PAP therapy is warranted in those requiring further respiratory support. If surgical management is required, tongue-lip adhesion or mandibular distraction osteogenesis have been shown to improve the AHI and relieve upper airway obstruction [111]. Because upper airway dimensions can improve over time [112], PAP therapy is a good alternative to consider prior to mandibular surgeries.

Neurologic Disorders

Children with neurologic disorders can be affected by a number of sleep abnormalities including OSAS, nocturnal obstructive hypoventilation, and diurnal hypoventilation. Pre-disposing factors to OSAS include reduced ventilatory responses, reduced activity of respiratory muscles during sleep, reduced upper airway dilator tone, and poor lung mechanics due to the underlying neuromuscular disorder [113]. Depending on the severity of disease, PAP therapy can be accomplished with CPAP, BPAP, or invasive ventilation and tracheotomy may be required in severe cases. For further discussion of sleep in children with neurologic disorders the reader is directed elsewhere [113].

Conclusion

In the coming years, pediatric OSAS will continue to burden our society. Improved methodologies to better diagnose and manage children with this disorder will continue to be developed as we improve our knowledge of its pathophysiology and the mechanisms leading to its adverse outcomes.

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Compliance with Ethical Standards

Conflict of interest The authors declare that they have no conflicts of interest.

References

- Marcus CL, Brooks LJ, Draper KA, Gozal D, Halbower AC, Jones J, Schechter MS, Sheldon SH, Spruyt K, Ward SD, Lehmann C, Shiffman RN (2012) Diagnosis and management of childhood obstructive sleep apnea syndrome. *Pediatrics* 130:576–584. <https://doi.org/10.1542/peds.2012-1671>
- Muzumdar H, Arens R (2013) Physiological effects of obstructive sleep apnea syndrome in childhood. *Respir Physiol Neurobiol* 188:370–382. <https://doi.org/10.1016/j.resp.2013.05.006>
- Lumeng JC, Chervin RD (2008) Epidemiology of pediatric obstructive sleep apnea. *Proc Am Thorac Soc* 5:242–252. <https://doi.org/10.1513/pats.200708-135mg>
- Arens R, Muzumdar H (2010) Childhood obesity and obstructive sleep apnea syndrome. *J Appl Physiol* 108:436–444. <https://doi.org/10.1152/japplphysiol.00689.2009>
- Tan H-L, Gozal D, Kheirandish-Gozal L (2013) Obstructive sleep apnea in children: a critical update. *Nat Sci Sleep*. <https://doi.org/10.2147/nss.s51907>
- Smith DF, Amin RS (2019) OSA and cardiovascular risk in pediatrics. *Chest* 156:402–413. <https://doi.org/10.1016/j.chest.2019.02.011>
- Marcus CL, Moore RH, Rosen CL, Giordani B, Garetz SL, Taylor HG, Mitchell RB, Amin R, Katz ES, Arens R, Paruthi S, Muzumdar H, Gozal D, Thomas NH, Janice Ware DB, Snyder K, Elden L, Sprecher RC, Willging P, Jones D, Bent JP, Hoban T, Chervin RD, Ellenberg SS, Redline S (2013) A randomized trial of adenotonsillectomy for childhood sleep apnea. *N Engl J Med* 368:2366–2376. <https://doi.org/10.1056/NEJMoa1215881>
- Reuveni H, Simon T, Tal A, Elhayany A, Tarasiuk A (2002) Health care services utilization in children with obstructive sleep apnea syndrome. *Pediatrics* 110:68–72. <https://doi.org/10.1542/peds.110.1.68>
- Arens R, Marcus CL (2004) Pathophysiology of upper airway obstruction: A developmental perspective. *Sleep* 27:997–1019. <https://doi.org/10.1093/sleep/27.5.997>
- Mitchell RB, Archer SM, Ishman SL, Rosenfeld RM, Coles S, Finestone SA, Friedman NR, Giordano T, Hildrew DM, Kim TW, Lloyd RM, Parikh SR, Shulman ST, Walner DL, Walsh SA, Nnacheta LC (2019) Clinical Practice guideline: tonsillectomy in children (update). *Otolaryngol Neck Surg* 160:S1–S42. <https://doi.org/10.1177/0194599818801757>
- Friedman M, Wilson M, Lin HC, Chang HW (2009) Updated systematic review of tonsillectomy and adenoidectomy for treatment of pediatric obstructive sleep apnea/hypopnea syndrome. *Otolaryngol Head Neck Surg* 140:800–808. <https://doi.org/10.1016/j.otohns.2009.01.043>
- Bhattacharjee R, Kheirandish-Gozal L, Spruyt K, Mitchell RB, Promchiarak J, Simakajornboon N, Kaditis AG, Splaingard D, Splaingard M, Brooks LJ, Marcus CL, Sin S, Arens R, Verhulst SL, Gozal D (2010) Adenotonsillectomy outcomes in treatment of obstructive sleep apnea in children: a multicenter retrospective study. *Am J Respir Crit Care Med* 182:676–683. <https://doi.org/10.1164/rccm.200912-1930OC>
- Tauman R, Gulliver TE, Krishna J, Montgomery-Downs HE, O'Brien LM, Ivanenko A, Gozal D (2006) Persistence of obstructive sleep apnea syndrome in children after adenotonsillectomy. *J Pediatr* 149:803–808. <https://doi.org/10.1016/j.jpeds.2006.08.067>
- Carroll JL, McColley SA, Marcus CL, Curtis S, Loughlin GM (1995) Inability of clinical history to distinguish primary snoring from obstructive sleep apnea syndrome in children. *Chest* 108:610–618. <https://doi.org/10.1378/chest.108.3.610>
- Certal V, Catumbela E, Winck JC, Azevedo I, Teixeira-Pinto A, Costa-Pereira A (2012) Clinical assessment of pediatric obstructive sleep apnea: a systematic review and meta-analysis. *Laryngoscope* 122:2105–2114. <https://doi.org/10.1002/lary.23465>
- Gozal D, Wang M, Pope DW (2001) Objective sleepiness measures in pediatric obstructive sleep apnea. *Pediatrics* 108:693–697. <https://doi.org/10.1542/peds.108.3.693>
- Rosen CL (1999) Clinical features of obstructive sleep apnea hypoventilation syndrome in otherwise healthy children. *Pediatr Pulmonol* 27:403–409. [https://doi.org/10.1002/\(SICI\)1099-0496\(199906\)27:6<3c403:AID-PPUL7%3e3.0.CO;2-8](https://doi.org/10.1002/(SICI)1099-0496(199906)27:6<3c403:AID-PPUL7%3e3.0.CO;2-8)
- Perfect MM, Archbold K, Goodwin JL, Levine-Donnerstein D, Quan SF (2013) Risk of behavioral and adaptive functioning difficulties in youth with previous and current sleep disordered breathing. *Sleep*. <https://doi.org/10.5665/sleep.2536>
- O'Brien LM, Mervis CB, Holbrook CR, Bruner JL, Smith NH, McNally N, McCliment MC, Gozal D (2004) Neurobehavioral correlates of sleep-disordered breathing in children. *J Sleep Res* 13:165–172. <https://doi.org/10.1111/j.1365-2869.2004.00395.x>
- Hunter SJ, Gozal D, Smith DL, Philby MF, Kaylegian J, Kheirandish-Gozal L (2016) Effect of sleep-disordered breathing severity on cognitive performance measures in a large community cohort of young school-aged children. *Am J Respir Crit Care Med* 194:739–747. <https://doi.org/10.1164/rccm.201510-2099OC>

21. Brodsky L (1989) Modern assessment of tonsils and adenoids. *Pediatr Clin N Am* 36:1551–1569. [https://doi.org/10.1016/S0031-3955\(16\)36806-7](https://doi.org/10.1016/S0031-3955(16)36806-7)
22. Marcus CL, Greene MG, Carroll JL (1998) Blood pressure in children with obstructive sleep apnea. *Am J Respir Crit Care Med* 157:1098–1103. <https://doi.org/10.1164/ajrccm.157.4.9704080>
23. Amin RS, Carroll JL, Jeffries JL, Grone C, Bean JA, Chini B, Bokulic R, Daniels SR (2004) Twenty-four-hour ambulatory blood pressure in children with sleep-disordered breathing. *Am J Respir Crit Care Med* 169:950–956. <https://doi.org/10.1164/rccm.200309-1305oc>
24. Ingram DG, Singh AV, Ehsan Z, Birnbaum BF (2017) Obstructive sleep apnea and pulmonary hypertension in children. *Paediatr Respir Rev* 23:33–39. <https://doi.org/10.1016/j.prrv.2017.01.001>
25. Berry RB, Albertario CL, Harding SM (2018) The AASM manual for the scoring of sleep and associated events: rules, terminology and technical specifications, version 2.5. American Academy of Sleep Medicine, Darien
26. American Academy of Sleep Medicine (2014) International classification of sleep disorders, 3rd edn. American Academy of Sleep Medicine, Darien
27. Isaiah A, Hamdan H, Johnson RF, Naqvi K, Mitchell RB (2017) Very severe obstructive sleep apnea in children: outcomes of adenotonsillectomy and risk factors for persistence. *Otolaryngol Head Neck Surg (United States)* 157:128–134. <https://doi.org/10.1177/0194599817700370>
28. Roland PS, Rosenfeld RM, Brooks LJ, Friedman NR, Jones J, Kim TW, Kuhar S, Mitchell RB, Seidman MD, Sheldon SH, Jones S, Robertson P (2011) Clinical practice guideline: polysomnography for sleep-disordered breathing prior to tonsillectomy in children. *Otolaryngol Neck Surg* 145:S1–S15. <https://doi.org/10.1177/0194599811409837>
29. Brouillette R, Hanson D, David R, Klemka L, Anna S, Fernbach S, Hunt C (1984) A diagnostic approach to suspected obstructive sleep apnea in children. *J Pediatr* 105:10–14. [https://doi.org/10.1016/S0022-3476\(84\)80348-0](https://doi.org/10.1016/S0022-3476(84)80348-0)
30. Chervin RD, Weatherly RA, Garetz SL, Ruzicka DL, Giordani BJ, Hodges EK, Dillon JE, Guire KE (2007) Pediatric sleep questionnaire: prediction of sleep apnea and outcomes. *Arch Otolaryngol Head Neck Surg* 133:216–222. <https://doi.org/10.1001/archotol.133.3.216>
31. Franco RA, Rosenfeld RM, Rao M (2000) Quality of life for children with obstructive sleep apnea. *Otolaryngol Neck Surg* 123:9–16. <https://doi.org/10.1067/mhn.2000.105254>
32. Patel AP, Meghji S, Phillips JS (2019) Accuracy of clinical scoring tools for the diagnosis of pediatric obstructive sleep apnea. *Laryngoscope*. <https://doi.org/10.1002/lary.28146>
33. Rosen CL, Wang R, Taylor HG, Marcus CL, Katz ES, Paruthi S, Arens R, Muzumdar H, Garetz SL, Mitchell RB, Jones D, Weng J, Ellenberg S, Redline S, Chervin RD (2015) Utility of symptoms to predict treatment outcomes in obstructive sleep apnea syndrome. *Pediatrics* 135:e662–e671. <https://doi.org/10.1542/peds.2014-3099>
34. Villa MP, Paolino MC, Castaldo R, Vanacore N, Rizzoli A, Miano S, Del Pozzo M, Montesano M (2013) Sleep clinical record: an aid to rapid and accurate diagnosis of paediatric sleep disordered breathing. *Eur Respir J* 41:1355–1361. <https://doi.org/10.1183/09031936.00215411>
35. Certal V, Silva H, Carvalho C, Costa-Pereira A, Azevedo I, Winck J, Capasso R, Camacho M (2015) Model for prediction of pediatric OSA: proposal for a clinical decision rule. *Laryngoscope* 125:2823–2827. <https://doi.org/10.1002/lary.25438>
36. Kapur VK, Auckley DH, Chowdhuri S, Kuhlmann DC, Mehra R, Ramar K, Harrod CG (2017) Clinical practice guideline OSA american academy. *J Clin Sleep Med* 13:479–504. <https://doi.org/10.5664/jcs.6506>
37. Kirk V, Baughn J, D'Andrea L, Friedman N, Galion A, Garetz S, Hassan F, Wrede J, Harrod CG, Malhotra RK (2017) American academy of sleep medicine position paper for the use of a home sleep apnea test for the diagnosis of OSA in children. *J Clin Sleep Med* 13:1199–1203. <https://doi.org/10.5664/jcs.6772>
38. Brouillette RT, Morielli A, Leimanis A, Waters KA, Luciano R, Ducharme FM (2000) Nocturnal pulse oximetry as an abbreviated testing modality for pediatric obstructive sleep apnea. *Pediatrics* 105:405–412. <https://doi.org/10.1542/peds.105.2.405>
39. Owen GO, Canter RJ (1996) Overnight pulse oximetry in normal children and in children undergoing adenotonsillectomy. *Clin Otolaryngol Allied Sci* 21:59–65. <https://doi.org/10.1111/j.1365-2273.1996.tb01026.x>
40. Hornero R, Kheirandish-Gozal L, Gutiérrez-Tobal GC, Philby MF, Alonso-Álvarez ML, Alvarez D, Dayyat EA, Xu Z, Huang YS, Kakazu MT, Li AM, Van Eyck A, Brockmann PE, Ehsan Z, Simakajornboon N, Kaditis AG, Vaquerizo-Villar F, Sedano AC, Capdevila OS, Von Lukowicz M, Terán-Santos J, Del CF, Poets CF, Ferreira R, Bertran K, Zhang Y, Schuen J, Verhulst S, Gozal D (2017) Nocturnal oximetry-based evaluation of habitually snoring children. *Am J Respir Crit Care Med* 196:1591–1598. <https://doi.org/10.1164/rccm.201705-0930OC>
41. Fujioka M, Young LW, Girdany BR (1979) Radiographic evaluation of adenoidal size in children: adenoidal-nasopharyngeal ratio. *Am J Roentgenol* 133:401–404. <https://doi.org/10.2214/ajr.133.3.401>
42. Kawashima S, Niikuni N, Chia-hung L, Takahasi Y, Kohno M, Nakajima I, Akasaka M, Sakata H, Akashi S (2000) Cephalometric comparisons of craniofacial and upper airway structures in young children with obstructive sleep apnea syndrome. *Ear Nose Throat J* 79:499–502. <https://doi.org/10.1177/014556130007900708>
43. Arens R, McDonough JM, Costarino AT, Mahboubi S, Tayag-Kier CE, Maislin G, Schwab RJ, Pack AI (2001) Magnetic resonance imaging of the upper airway structure of children with obstructive sleep apnea syndrome. *Am J Respir Crit Care Med* 164:698–703. <https://doi.org/10.1164/ajrccm.164.4.2101127>
44. Gibson SE, Strife JL, Myer CM, O'Connor DM (1996) Sleep fluoroscopy for localization of upper airway obstruction in children. *Ann Otol Rhinol Laryngol* 105:678–683. <https://doi.org/10.1177/000348949610500902>
45. Slaats MA, Van Hoorenbeeck K, Van Eyck A, Vos WG, De Backer JW, Boudewyns A, De Backer W, Verhulst SL (2015) Upper airway imaging in pediatric obstructive sleep apnea syndrome. *Sleep Med Rev* 21:59–71. <https://doi.org/10.1016/j.smrv.2014.08.001>
46. Fregosi RF, Quan SF, Kaemingk KL, Morgan WJ, Goodwin JL, Cabrera R, Gmitro A (2003) Sleep-disordered breathing, pharyngeal size and soft tissue anatomy in children. *J Appl Physiol* 95:2030–2038. <https://doi.org/10.1152/japplphysiol.00293.2003>
47. Arens R, Sin S, McDonough JM, Palmer JM, Dominguez T, Meyer H, Wootton DM, Pack AI (2005) Changes in upper airway size during tidal breathing in children with obstructive sleep apnea syndrome. *Am J Respir Crit Care Med* 171:1298–1304. <https://doi.org/10.1164/rccm.200411-1597OC>
48. Wootton DM, Sin S, Luo H, Yazdani A, McDonough JM, Wagshul ME, Isasi CR, Arens R (2016) Computational fluid dynamics upper airway effective compliance, critical closing pressure, and obstructive sleep apnea severity in obese adolescent girls. *J Appl Physiol* 121:925–931. <https://doi.org/10.1152/japplphysiol.00190.2016>
49. Shott SR, Donnelly LF (2004) Cine magnetic resonance imaging: Evaluation of persistent airway obstruction after tonsil and

- adenoidectomy in children with down syndrome. *Laryngoscope* 114:1724–1729. <https://doi.org/10.1097/00005537-200410000-00009>
50. Socarras MA, Landau BP, Durr ML (2019) Diagnostic techniques and surgical outcomes for persistent pediatric obstructive sleep apnea after adenotonsillectomy: A systematic review and meta-analysis. *Int J Pediatr Otorhinolaryngol* 121:179–187. <https://doi.org/10.1016/j.ijporl.2019.02.030>
 51. Wilcox LJ, Bergeron M, Reghunathan S, Ishman SL (2017) An updated review of pediatric drug-induced sleep endoscopy. *Laryngosc Investig Otolaryngol* 2:423–431. <https://doi.org/10.1002/liv.118>
 52. He S, Peddireddy NS, Smith DF, Duggins AL, Heubi C, Shott SR, Ishman SL (2018) Outcomes of drug-induced sleep endoscopy-directed surgery for pediatric obstructive sleep apnea. *Otolaryngol Head Neck Surg* 158:559–565. <https://doi.org/10.1177/0194599817740332>
 53. Esteller E, Villatoro JC, Agüero A, Matión E, Lopez R, Arístimuño A, Nuñez V, Díaz-Herrera MA (2019) Outcome of drug-induced sleep endoscopy-directed surgery for persistent obstructive sleep apnea after adenotonsillar surgery. *Int J Pediatr Otorhinolaryngol* 120:118–122. <https://doi.org/10.1016/j.ijporl.2019.02.004>
 54. Drager LF, Togeiro SM, Polotsky VY, Lorenzi-Filho G (2013) Obstructive sleep apnea: a cardiometabolic risk in obesity and the metabolic syndrome. *J Am Coll Cardiol* 62:569–576. <https://doi.org/10.1016/j.jacc.2013.05.045>
 55. Chinnadurai S, Jordan AK, Sathe NA, Fannesbeck C, McPheeters ML, Francis DO (2017) Tonsillectomy for obstructive sleep-disordered breathing: a meta-analysis. *Pediatrics* 139:e20163491. <https://doi.org/10.1542/peds.2016-3491>
 56. Sulman CG (2014) Pediatric sleep surgery. *Front Pediatr* 2:51. <https://doi.org/10.3389/fped.2014.00051>
 57. Parikh SR, Archer S, Ishman SL, Mitchell RB (2019) why is there no statement regarding partial intracapsular tonsillectomy (tonsillotomy) in the new guidelines? *Otolaryngol Head Neck Surg* 160:213–214. <https://doi.org/10.1177/0194599818810507>
 58. Mukhatiyar P, Nandalike K, Cohen HW, Sin S, Gangar M, Bent JP, Arens R (2016) Intracapsular and extracapsular tonsillectomy and adenoidectomy in pediatric obstructive sleep apnea. *JAMA Otolaryngol Head Neck Surg* 142:25–31. <https://doi.org/10.1001/jamaoto.2015.2603>
 59. Imanguli M, Ulualp SO (2016) Risk factors for residual obstructive sleep apnea after adenotonsillectomy in children. *Laryngoscope* 126:2624–2629. <https://doi.org/10.1002/lary.25979>
 60. Bluher AE, Ishman SL, Baldassari CM (2019) Managing the child with persistent sleep apnea. *Otolaryngol Clin N Am* 52:891–901. <https://doi.org/10.1016/j.otc.2019.06.004>
 61. Com G, Carroll JL, Tang X, Melguizo MS, Bower C, Jambhekar S (2015) Characteristics and surgical and clinical outcomes of severely obese children with obstructive sleep apnea. *J Clin Sleep Med* 11:467–474. <https://doi.org/10.5664/jcsm.4608>
 62. Ulualp SO (2014) Modified expansion sphincter pharyngoplasty for treatment of children with obstructive sleep apnea. *JAMA Otolaryngol Head Neck Surg* 140:817–822. <https://doi.org/10.1001/jamaoto.2014.1329>
 63. Cheng PW, Fang KM, Su HW, Huang TW (2012) Improved objective outcomes and quality of life after adenotonsillectomy with inferior turbinate reduction in pediatric obstructive sleep apnea with inferior turbinate hypertrophy. *Laryngoscope* 122:2850–2854. <https://doi.org/10.1002/lary.23590>
 64. Grindle CR, Murray RC, Chennupati SK, Barth PC, Reilly JS (2011) Incidence of revision adenoidectomy in children. *Laryngoscope* 121:2128–2130. <https://doi.org/10.1002/lary.22161>
 65. Noller MW, Guillemineault C, Gouveia CJ, Mack D, Neighbors CL, Zaghi S, Camacho M (2018) Mandibular advancement for pediatric obstructive sleep apnea: a systematic review and meta-analysis. *J Cranio-Maxillofacial Surg* 46:1296–1302. <https://doi.org/10.1016/j.jcms.2018.04.027>
 66. Tahiri Y, Viezel-Mathieu A, Aldekhayel S, Lee J, Gilardino M (2014) The effectiveness of mandibular distraction in improving airway obstruction in the pediatric population. *Plast Reconstr Surg* 133:352–359. <https://doi.org/10.1097/01.prs.0000438049.29258.a8>
 67. Wootten CT, Shott SR (2010) Evolving therapies to treat retroglossal and base-of-tongue obstruction in pediatric obstructive sleep apnea. *Arch Otolaryngol Head Neck Surg* 136:983–987. <https://doi.org/10.1001/archoto.2010.178>
 68. Song SA, Wei JM, Buttram J, Tolisano AM, Chang ET, Liu SYC, Certal V, Camacho M (2016) Hyoid surgery alone for obstructive sleep apnea: a systematic review and meta-analysis. *Laryngoscope* 126:1702–1708. <https://doi.org/10.1002/lary.25847>
 69. Rizzi CJ, Amin JD, Isaiah A, Valdez TA, Jeyakumar A, Smart SE, Pereira KD (2017) Tracheostomy for severe pediatric obstructive sleep apnea: indications and outcomes. *Otolaryngol Head Neck Surg (United States)* 157:309–313. <https://doi.org/10.1177/0194599817702369>
 70. Berry RB, Chediak A, Brown LK, Finder J, Gozal D, Iber C, Kushida CA, Morgenthaler T, Rowley JA, Davidson-Ward SL (2010) Best clinical practices for the sleep center adjustment of noninvasive positive pressure ventilation (NPPV) in stable chronic alveolar hypoventilation syndromes: NPPV titration task force of the American academy of sleep medicine. *J Clin Sleep Med* 6:491–509
 71. Marcus CL, Rosen G, Ward SLD, Halbower AC, Sterni L, Lutz J, Stading PJ, Bolduc D, Gordon N (2006) Adherence to and effectiveness of positive airway pressure therapy in children with obstructive sleep apnea. *Pediatrics* 117:e442–e451. <https://doi.org/10.1542/peds.2005-1634>
 72. Uong EC, Epperson M, Bathon SA, Jeffe DB (2007) Adherence to nasal positive airway pressure therapy among school-aged children and adolescents with obstructive sleep apnea syndrome. *Pediatrics* 120:e1203–e1211. <https://doi.org/10.1542/peds.2006-2731>
 73. Haentjens P, Van Meerhaeghe A, Moscariello A, De Weerd S, Poppe K, Dupont A, Velkeniers B (2007) The impact of continuous positive airway pressure on blood pressure in patients with obstructive sleep apnea syndrome: evidence from a meta-analysis of placebo-controlled randomized trials. *Arch Intern Med* 167:757–765. <https://doi.org/10.1001/archinte.167.8.757>
 74. Giles TL, Lasserson TJ, Smith BH, White J, Wright J, Cates CJ (2006) Continuous positive airways pressure for obstructive sleep apnoea in adults. *Cochrane database Syst Rev*. <https://doi.org/10.1002/14651858.CD001106.pub3>
 75. Jambhekar SK, Com G, Tang X, Pruss KK, Jackson R, Bower C, Carroll JL, Ward W (2013) Role of a respiratory therapist in improving adherence to positive airway pressure treatment in a pediatric sleep apnea clinic. *Respir Care* 58:2038–2044. <https://doi.org/10.4187/respcare.02312>
 76. Kirk VG, O'Donnell AR (2006) Continuous positive airway pressure for children: a discussion on how to maximize compliance. *Sleep Med Rev* 10:119–127. <https://doi.org/10.1016/j.smrv.2005.07.002>
 77. Pépin JL, Leger P, Veale D, Langevin B, Robert D, Lévy P (1995) Side effects of nasal continuous positive airway pressure in sleep apnea syndrome. Study of 193 patients in two French sleep centers. *Chest* 107:375–381. <https://doi.org/10.1378/chest.107.2.375>
 78. Ahmad Z, Venus M, Kisku W, Rayatt SS (2013) A case series of skin necrosis following use of non invasive ventilation pressure masks. *Int Wound J* 10:87–90. <https://doi.org/10.1111/j.1742-481X.2012.00948.x>

79. Li KK, Riley RW, Guilleminault C (2000) An unreported risk in the use of home nasal continuous positive airway pressure and home nasal ventilation in children: mid-face hypoplasia. *Chest* 117:916–918. <https://doi.org/10.1378/chest.117.3.916>
80. Roberts SD, Kapadia H, Greenlee G, Chen ML (2016) Midfacial and dental changes associated with nasal positive airway pressure in children with obstructive sleep apnea and craniofacial conditions. *J Clin Sleep Med* 12:469–475. <https://doi.org/10.5664/jcsm.5668>
81. Fauroux B, Lavis J-F, Nicot F, Picard A, Boelle P-Y, Clément A, Vazquez M-P (2005) Facial side effects during noninvasive positive pressure ventilation in children. *Intens Care Med* 31:965–969. <https://doi.org/10.1007/s00134-005-2669-2>
82. Gudnadottir G, Ellegård E, Hellgren J (2018) Intranasal budesonide and quality of life in pediatric sleep-disordered breathing: a randomized controlled trial. *Otolaryngol Head Neck Surg (United States)* 158:752–759. <https://doi.org/10.1177/0194599817742597>
83. Kheirandish L, Goldbart AD, Gozal D (2006) Intranasal steroids and oral leukotriene modifier therapy in residual sleep-disordered breathing after tonsillectomy and adenoidectomy in children. *Pediatrics*. <https://doi.org/10.1542/peds.2005-0795>
84. Liming BJ, Ryan M, Mack D, Ahmad I, Camacho M (2019) Montelukast and nasal corticosteroids to treat pediatric obstructive sleep apnea: a systematic review and meta-analysis. *Otolaryngol Head Neck Surg (United States)* 160:594–602. <https://doi.org/10.1177/0194599818815683>
85. Kheirandish-Gozal L, Bhattacharjee R, Bandla HPR, Gozal D (2014) Antiinflammatory therapy outcomes for mild OSA in children. *Chest* 146:88–95. <https://doi.org/10.1378/chest.13-2288>
86. Villa MP, Malagola C, Pagani J, Montesano M, Rizzoli A, Guilleminault C, Ronchetti R (2007) Rapid maxillary expansion in children with obstructive sleep apnea syndrome: 12-month follow-up. *Sleep Med* 8:128–134. <https://doi.org/10.1016/j.sleep.2006.06.009>
87. Pirelli P, Saponara M, Guilleminault C (2004) Rapid maxillary expansion in children with obstructive sleep apnea syndrome. *Sleep* 27:761–766. <https://doi.org/10.1093/sleep/27.4.761>
88. Villa MP, Bernkopf E, Pagani J, Broia V, Montesano M, Ronchetti R (2002) Randomized controlled study of an oral jaw-positioning appliance for the treatment of obstructive sleep apnea in children with malocclusion. *Am J Respir Crit Care Med* 165:123–127. <https://doi.org/10.1164/ajrccm.165.1.2011031>
89. Carvalho FR, Lentini-Oliveira DA, Prado LB, Prado GF, Carvalho LBC (2016) Oral appliances and functional orthopaedic appliances for obstructive sleep apnoea in children. *Cochrane database Syst Rev*. <https://doi.org/10.1002/14651858.CD005520.pub3>
90. Shete MM, Stocks RMS, Sebelik ME, Schoumacher RA (2010) Effects of adeno-tonsillectomy on polysomnography patterns in down syndrome children with obstructive sleep apnea: a comparative study with children without Down syndrome. *Int J Pediatr Otorhinolaryngol* 74:241–244. <https://doi.org/10.1016/j.ijporl.2009.11.006>
91. Redline S, Tishler PV, Schluchter M, Aylor J, Clark K, Graham G (1999) Risk factors for sleep-disordered breathing in children: associations with obesity, race, and respiratory problems. *Am J Respir Crit Care Med* 159:1527–1532. <https://doi.org/10.1164/ajrccm.159.5.9809079>
92. Arens R, Sin S, Nandalike K, Rieder J, Khan UI, Freeman K, Wylie-Rosett J, Lipton ML, Wootton DM, McDonough JM, Shifteh K (2011) Upper airway structure and body fat composition in obese children with obstructive sleep apnea syndrome. *Am J Respir Crit Care Med* 183:782–787. <https://doi.org/10.1164/rccm.201008-1249OC>
93. Andersen IG, Holm JC, Homøe P (2016) Obstructive sleep apnea in obese children and adolescents, treatment methods and outcome of treatment - A systematic review. *Int J Pediatr Otorhinolaryngol* 87:190–197. <https://doi.org/10.1016/j.ijporl.2016.06.017>
94. Nandalike K, Shifteh K, Sin S, Strauss T, Stakofsky A, Gonik N, Bent J, Parikh SR, Bassila M, Nikova M, Muzumdar H, Arens R (2013) Adenotonsillectomy in obese children with obstructive sleep apnea syndrome: magnetic resonance imaging findings and considerations. *Sleep* 36:841–847. <https://doi.org/10.5665/sleep.2708>
95. Andersen IG, Holm JC, Homøe P (2019) Impact of weight-loss management on children and adolescents with obesity and obstructive sleep apnea. *Int J Pediatr Otorhinolaryngol* 123:57–62. <https://doi.org/10.1016/j.ijporl.2019.04.031>
96. Styne DM, Arslanian SA, Connor EL, Farooqi IS, Murad MH, Silverstein JH, Yanovski JA (2017) Pediatric obesity-assessment, treatment, and prevention: an endocrine society clinical practice guideline. *J Clin Endocrinol Metab* 102:709–757. <https://doi.org/10.1210/jc.2016-2573>
97. Scheffler P, Wolter NE, Narang I, Amin R, Holler T, Ishman SL, Propst EJ (2019) Surgery for obstructive sleep apnea in obese children: literature review and meta-analysis. *Otolaryngol Head Neck Surg (United States)* 160:985–992. <https://doi.org/10.1177/0194599819829415>
98. Lee CF, Lee CH, Hsueh WY, Lin MT, Kang KT (2018) Prevalence of obstructive sleep apnea in children with down syndrome: a meta-analysis. *J Clin Sleep Med* 14:867–875. <https://doi.org/10.5664/jcsm.7126>
99. Bull MJ, Saal HM, Braddock SR, Enns GM, Gruen JR, Perrin JM, Saul RA, Tarini BA, Hersh JH, Mendelsohn NJ, Hanson JW, Lloyd-Puryear MA, Musci TJ, Rasmussen SA, Downs SM, Spire P (2011) Clinical report—health supervision for children with down syndrome. *Pediatrics* 128:393–406. <https://doi.org/10.1542/peds.2011-1605>
100. Uong EC, McDonough JM, Tayag-Kier CE, Zhao H, Hasselgrove J, Mahboubi S, Schwab RJ, Pack AI, Arens R (2001) Magnetic resonance imaging of the upper airway in children with down syndrome. *Am J Respir Crit Care Med* 163:731–736. <https://doi.org/10.1164/ajrccm.163.3.2004231>
101. Fricke BL, Donnelly LF, Shott SR, Kalra M, Poe SA, Chini BA, Amin RS (2006) Comparison of lingual tonsil size as depicted on MR imaging between children with obstructive sleep apnea despite previous tonsillectomy and adenoidectomy and normal controls. *Pediatr Radiol* 36:518–523. <https://doi.org/10.1007/s00247-006-0149-7>
102. Donnelly LF, Shott SR, LaRose CR, Chini BA, Amin RS (2004) Causes of persistent obstructive sleep apnea despite previous tonsillectomy and adenoidectomy in children with down syndrome as depicted on static and dynamic cine MRI. *Am J Roentgenol* 183:175–181. <https://doi.org/10.2214/ajr.183.1.1830175>
103. Hawkins A, Langton-Hewer S, Henderson J, Tulloh RM (2011) Management of pulmonary hypertension in down syndrome. *Eur J Pediatr* 170:915–921. <https://doi.org/10.1007/s00431-010-1378-1>
104. Carskadon MA, Puschel SM, Millman RP (1993) Sleep-disordered breathing and behavior in three risk groups: preliminary findings from parental reports. *Child's Nerv Syst* 9:452–457. <https://doi.org/10.1007/BF00393547>

105. Breslin J, Spanò G, Bootzin R, Anand P, Nadel L, Edgin J (2014) Obstructive sleep apnea syndrome and cognition in down syndrome. *Dev Med Child Neurol* 56:657–664. <https://doi.org/10.1111/dmcn.12376>
106. Lal C, White DR, Joseph JE, Van Bakergem K, LaRosa A (2015) Sleep-disordered breathing in down syndrome. *Chest* 147:570–579. <https://doi.org/10.1378/chest.14-0266>
107. Farhood Z, Isley JW, Ong AA, Nguyen SA, Camilon TJ, LaRosa AC, White DR (2017) Adenotonsillectomy outcomes in patients with Down syndrome and obstructive sleep apnea. *Laryngoscope* 127:1465–1470. <https://doi.org/10.1002/lary.26398>
108. Akkina SR, Ma CC, Kirkham EM, Horn DL, Chen ML, Parikh SR (2018) Does drug induced sleep endoscopy-directed surgery improve polysomnography measures in children with down Syndrome and obstructive sleep apnea? *Acta Otolaryngol* 138:1009–1013. <https://doi.org/10.1080/00016489.2018.1504169>
109. Anderson ICW, Sedaghat AR, McGinley BM, Redett RJ, Boss EF, Ishman SL (2011) Prevalence and severity of obstructive sleep apnea and snoring in infants with Pierre Robin sequence. *Cleft Palate-Craniofacial J* 48:614–618. <https://doi.org/10.1597/10-100>
110. Kurian C, Ehsan Z (2019) Sleep and respiratory outcomes in neonates with Pierre Robin sequence: a concise review. *Sleep Breath*. <https://doi.org/10.1007/s11325-019-01876-6>
111. Hsieh ST, Woo AS (2019) Pierre Robin sequence. *Clin Plast Surg* 46:249–259. <https://doi.org/10.1016/j.cps.2018.11.010>
112. Staudt CB, Gnoinski WM, Peltomäki T (2013) Upper airway changes in Pierre Robin sequence from childhood to adulthood. *Orthod Craniofac Res* 16:202–213. <https://doi.org/10.1111/ocr.12019>
113. Arens R, Muzumdar H (2010) Sleep, sleep disordered breathing, and nocturnal hypoventilation in children with neuromuscular diseases. *Paediatr Respir Rev* 11:24–30. <https://doi.org/10.1016/j.prrv.2009.10.003>
114. Aurora RN, Zak RS, Karippot A, Lamm CI, Morgenthaler TI, Auerbach SH, Bista SR, Casey KR, Chowdhuri S, Kristo DA, Ramar K (2011) Practice parameters for the respiratory indications for polysomnography in children. *Sleep* 34:379–388. <https://doi.org/10.1093/sleep/34.3.379>
115. Kaditis AG, Alvarez MLA, Boudewyns A, Alexopoulos EI, Ersu R, Joosten K, Larramona H, Miano S, Narang I, Trang H, Tsaousoglou M, Vandenbussche N, Villa MP, Van WD, Weber S, Verhulst S (2016) Obstructive sleep disordered breathing in 2- to 18-year-old children: diagnosis and management. *Eur Respir J* 47:69–94. <https://doi.org/10.1183/13993003.00385-2015>

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