Understanding Nasal Breathing: the key to evaluating and treating sleep disordered breathing in adults and children

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Abstract

Nasal breathing is a basic and critical function that we rarely think much about, and most would be quite surprised to learn how important and influential it is in growth and development, and a healthy productive life. Fundamentally speaking, nasal breathing provides us with a sense of smell, enhances oxygen absorption by our lungs (via nitric oxide production from the sinuses, warms and humidifies the air we breathe before it reaches the lower airway, and helps filter impurities from the air. However, it is also responsible for proper craniofacial development' temporo-mandibular joint function head posture and more. When nasal breathing does not occur for any reason, the body is forced into a series of compromises that prioritize getting oxygen into our blood at the expense of the functions provided by the nasal breathing. Because oxygen is required for survival, we are forced to live with a host of acquired health issues as a consequence of chronic or intermittent mouth breathing. One such consequence, known as sleep disordered breathing (SDB), is becoming pandemic in children, and directly affects their growth, development, intellect, academic performance, behavior, and much more. In the article, we will review the causes of nasal obstruction, and how it relates to craniofacial development and palatal growth, the development of ADD and ADHD, and new minimally invasive dental, nasal and airway procedures to correct the problem. On this review we will assess the different symptoms and causes, as palate shape changes or tonsils disease. Also a quick view about available therapeutic approaches will be done, including minimal invasive nasal surgery as well support devices as linguaflex an CPAP.

<u>The Common Etiologies of Nasal</u> <u>Obstruction</u>

breathing Nasal is an extremely important function, and we have come to appreciate that many health problems^{1,2,3,4,5} develop when nasal breathing is compromised. The etiology of reduced nasal airflow is limited, and includes the following 5 major categories. Anatomic deformities, such as a septal deviation, alar collapse, turbinate hypertrophy, or the presence of concha bullosa are relatively common. These problems are typically genetic, but the first two can also be the result of nasal trauma. Atopy can cause swelling of the nasal submucosa and mucosal glands, and thus limit airflow. The majority of glands are located in the nasal turbinates and septal swell body, and are a major cause of their hypertrophy.¹ Atopy can be treated with various medications, however, while topical nasal steroids are the preferred treatment, a paradox arises when nasal congestion actually minimizes the access of topical medications to the nose, this markedly decreasing their effectiveness. A nasal mass or nasal polyps can also cause nasal obstruction, with symptoms that can include congestion, bloody discharge, facial pain or pressure, headache, and hyposmia. These growths can occur in children and adults, be benign or malignant, and have the best prognosis when diagnosed and treated Adenoid hypertrophy is very early. common in children and can cause severe nasal obstruction. The adenoids typically shrink by age > 4. Adenoids, like tonsils, are part of our immune system and serve little purpose after age 2^7 . In addition to nasal obstruction, enlarged adenoids are implicated in the development of chronic serous otitis media and chronic sinusitis in children⁸. Vasomotor rhinitis (VMR) is perhaps the challenging most cause of nasal obstruction and is due to and imbalance in the autonomic nervous system which controls nasal blood flow. This leads to intermittent vascular engorgement on one side or in one part of the nose, and is occasionally non-remitting. Improved breathing may occur sporadically during the day, and symptoms are almost always worse after lying down, especially on one's side. As with most conditions, there are varying degrees of VMR. These patients often describe alternating nasal congestion, especially when supine. It is also possible to have several of these factors present simultaneously (i.e. anatomic and atopic, or atopy and polyps).

Symptoms Differ in Adults vs Children

The most common etiology of nasal obstruction differs between adults and children. For the former they are, in order of decreasing frequency, anatomic deformities, allergy, VMR, a nasal mass (adenoid remnants are extremely rare); whereas in the latter the order would be adenoid hypertrophy. anatomic deformities, allergy, VMR, and a nasal mass. Regardless of the cause(s), nasal obstruction in adults and children leads to major physiologic changes that can have serious health consequences. In adults, a common example of the consequence of nasal obstruction is snoring and obstructive sleep apnea (OSA). The most common symptom of OSA in adults is daytime fatigue and cognitive impairment, followed by a significantly increased risk for heart disease, high blood pressure, stroke,

diabetes, memory loss, and reduced libido⁹. Bruxism and TMJ dysfunction are now also linked to nasal obstruction in sleep³. In children, OSA and sleep disordered breathing (SDB) leads to very different signs and symptoms that are very common, yet often overlooked or misunderstood. These include mouth breathing, snoring, restless sleep, frequent awakenings, enuresis, lack of head-forward focus. posturing, disinterest in eating, attention deficit disorder (ADD), hyperactivity (ADHD), poor school performance, anger, and frank aggression¹⁰. Headaches may also be a significant complaint. Teens can show more of the daytime fatigue seen in adults.

A key point to remember is that not all symptoms are present in each patient and the exact constellation of symptoms varies widely among patients. For patients with OSA and SDB, more sleep is not helpful because it is during sleep when their heart, brain and other organs are stressed due to oxygen debt. They are also in what is called "*sympathetic dystrophy*", where their body is subject to continuous release of adrenaline during sleep. The repetitive adrenaline surge (often called arousals on a polysomnogram) has very different physiologic effects in adults versus children. More on this to follow. CPAP (continuous positive airway pressure) machines, worn on the face or nose, attempt to force oxygen past the nasal obstruction to provide adequate oxygen during sleep. (Fig 1) However, CPAP tolerance in generally quite poor, with large sample data showing few patients using it more than 4 hours a night¹¹.

Fig 1, CPAP devices in use





CPAP

Another very important point is that many patients with OSA or SDB are unaware of their nasal congestion. The two most common reasons for this are that in many patients, nasal obstruction is positional and only occurs once the patient is lying down for sleep. It takes anywhere from 30 to 90 minutes for the nasal congestion to develop when supine, and by then the patient is often already asleep and unaware of the problem. The second reason is the variability in subjective appreciation or self-awareness of nasal compromise. Many of these patients have had nasal obstruction since early childhood, thus they have no frame of reference for normal nasal breathing. It is for this very reason that we test children for hearing and visual loss in grade school. Children have no frame of reference for normal hearing or sight. Likewise, they have no frame of reference for normal nasal breathing, and unfortunately, there is no standard screening test for them at this time. Another common compensation for nasal obstruction is forward head posturing¹⁰. In order to mouth breath better, the oral airway needs to be aligned with the trachea, this would produce a chin up, extended head position. (Fig 2) Many children will sleep in an extended head position for

this reason. (Fig 3) However, during daytime, we cannot see where we are going in this position. Therefore, we must lean our extended neck forward. This forward head posturing places significant stress on the posterior neck muscles and leads to cervical spine alignment. Poor cervical spine alignment can lead to nerve root compression and mal-alignment of the lower spine. Thus, once the first domino falls, it is uncertain which will be the last domino to fall - i.e. will head forward posturing affect cervical spine position alone, or will it result in lower back problems, gait issues, neck pain, etc.

The bottom line is that many children who suffer from SDB go undiagnosed. Awareness for the signs and symptoms as described herein should alert teachers, parents, and physicians to the possibility of SDB in children.



Fig 3: Mouth breather with extended neck posture



The Primate Studies in Dentistry

That said, the most likely person to diagnose a child with SDB is their family dentist or orthodontist. This is directly related to the fact that nasal breathing has a tremendous impact on cranio-facial development¹², which in turn determines our dental profile and occlusion. Over 90% of children with crooked teeth, teeth grinding, or malocclusion have compromised nasal breathing¹³. (Fig 4, 5, 6) This important medical fact was first observed in the 1970's by a Norwegian orthodontist named Dr. Egil Harvold.

He noticed that many of his patients with these dental issues had nasal problems and were often mouth breathers. He reported his landmark study in 1981 entitled "Primate experiments in oral respiration"¹⁴, where he took two groups of baby monkeys and secured silicone plugs into the nostrils of half of them. He watched as the monkeys grew over the next 6 months and noticed the ones with nasal plugs began to mouth breath by default, which in turn lead to many changes in cranio-facial growth compared to the monkeys without nasal obstruction.



Fig 4. Craniofacial changes in mouth breathing young male



Fig 5. Craniofacial changes in mouth breathing young female

Fig 6. Craniofacial and dental changes in young female



He reported the narrow maxilla, high arched palate, elongated and downwards rotation of the mandible, and varying degrees of malocclusion were hallmarks of nasal obstruction in children. Not only were skeletal changes being observed, but compensatory changes by the masticator muscles were causing lower jaw malposition. The affected monkeys subsequently developed narrow faces with elongated mandibles, known today as "*adenoid facies*". (Fig 7) This finding is the physical hallmark of nasal obstruction in young children, and unfortunately, is an end-stage situation. Interestingly, Harvold then removed the nasal plugs from the monkeys who had them, and noted that many of the occlusion and jaw issues resolved over the next 6 months, therefore proving that restoration of nasal breathing can lead to reversal of these changes.





As a result of compromised nasal breathing, the narrow maxilla cannot accommodate the full complement of upper teeth and a crowded dental pattern develops. (Fig 8) It is important to note that if nasal obstruction is not corrected after the teeth have been repositioned, the once straightened teeth will likely shift again because the forces causing the shift has never been corrected. We frequently see adults and teens requiring a second, and sometimes third set of braces for this exact reason. In the past, orthodontists would extract teeth to "uncrowd" the maxilla and use braces to realign the remaining teeth. Historically, orthodontics has been the science of restorative aesthetic dentistry, whereas now it is far more focused on functional upper airway development. Thus, it is also important to diagnose and treat children at an early age because 60% of their facial growth occurs by age 4, 70% by age 5, and 90% by age 12^{15} . The old philosophy of waiting until a child is a teen or pre-teen to address their crooked teeth is no longer appropriate. Dental

extractions to un-crowd the maxilla are now largely abandoned in place of what is called palatal expansion¹⁶. Bv temporarily attaching an oral device onto the inner table of the alveolar ridge of the maxilla, the orthodontist can perform osteo-distraction, whereby they slowly move the palatal bones apart over a 6-12 month period. (Fig 9) There is no pain as the process is slow and gentle. Palatal expanders can actually bear the child's initials or name and have become an oral fashion statement for kids. Palatal expansion ultimately causes widening of the constricted maxilla and will allow proper alignment of all teeth using braces without requiring extractions.



Fig 9. Palatal expanders

1 mm of palatal expansion increases nasal airway 2.4% (Motro, et al, Boston Univ)



Palatal Expansion

It was recently reported by Dr. Melih Motro. researcher Boston a at Goldman University's School of Dentistry, that for every 1 mm of lateral palatal expansion, the nasal airway increases by $2.4\%^{17}$. This is an extremely important discovery because it shows us how a simple minimally invasive office treatment can begin to correct the craniofacial abnormalities that develop from reduced nasal breathing. Since the palatal bones in children are not yet fused in the midline,

it is relatively easy to obtain expansion of 4-10 mm in a relatively short time. Unfortunately, this is not as easy in adults because the palatal suture has already fused. As a result of this study, palatal expansion in children with SDB and dental mal-alignment is now standard practice as the first step in modern orthodontic care¹⁸. Wearing head-frames or other dental devices designed to correct these orthodontic problems are actually counter-productive and place restrictive forces on a maxilla that is developing in an anterior and inferior direction¹⁸.

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ADD/ADHD and Behavioral Changes in Children

It has also been shown with increasing evidence, that approximately 40% of children who suffer from SDB develop ADD, ADHD, and/or a learning disabilitv¹⁹. Statistically, if a child snores by the age of 8 and is untreated, there is an 80% chance the child will have a permanent 20% reduction in mental capacity²⁰. Additionally, if a child is diagnosed with SDB in the first 5 years of life and untreated, they are 60% more likely to require special needs education by age 8^{21} . However, the clinicians diagnosing these children and offering treatment for ADD/ADHD rarely consider nasal obstruction as a cause, and are also unaware that the condition is likely reversible without the need for medications or psychological therapy. ADD and ADHD are the direct result of the concept of sympathetic dystrophy I introduced earlier. Sleep deprived children develop hyperactive behavior, and this phenomenon is witnessed in most young children as their bedtime approaches. Exposing a child to hours of adrenaline over the course of a night leads to restless sleep at night and hours of hyperactive behavior the following day. It is also responsible for their enuresis, frequent awakenings, etc²². In adults, sympathetic dystrophy leads to hypertension, heart disease, stroke, and diabetes. The fatigue and cognitive issues are due to hours of oxygen debt during poor sleep. Note the vastly different phenotypes of SDB and OSA in children versus adults.

Our group is currently investigating the reversibility of ADD/ADHD in these children. Early results show a significant ability to reverse the condition once normal nasal breathing is established. While the results of this study are not complete, we are extremely encouraged by the early data and the significant number of children whose behavior has been positively affected by reversing mouth breathing tendencies and providing quality sleep. We are currently evaluating the data with respect to the child's age, gender, and severity of baseline ADD/ADHD to determine if there are any differences in with outcomes respect to these The ability to reverse or parameters.

reduce the affects of ADD/ADHD in children is a major breakthrough with profound quality of life, social, and family implications. We have even seen children with mild autism, anxiety, or aggression become highly functioning students to the astonishment of their parents and teachers. versus adults, and have introduced some treatment options. In the 1980's it was discovered that hypertrophic tonsils and adenoids were responsible for SDB and OSA, and at that time, it was considered the primary etiology²³. Hence, many children underwent T&A with the goal of correcting their SDB. Did we finally have a cure for SDB in children? (Fig 10)

<u>The Role of Tonsils & Adenoids</u>

Thus far, we have described the major causes of compromised nasal breathing, the different phenotypes in children

Fig 10. Images of large obstructing tonsils and adenoids



The answer is yes and no! Studies looking at large groups of children with SDB who underwent corrective T&A have shown that about 67% relapse with SDB > 2 years of surgery²⁴. This is a very important point that was revealed after longitudinal study of affected children, and shows the importance of long-term follow-up. It also highlights a knowledge gap between the dental and medical communities due to a lack of communication and knowledge sharing. I mentioned earlier how Dr. Harvold had discovered the importance of nasal obstruction in craniofacial development in 1981. Yet this information was not well known or shared with medical colleagues, and largely stayed within the dental community. T&A can improve nasal breathing and sleep in many patients, however, if we do not evaluate the other 4 causes of nasal obstruction mentioned earlier, we cannot expect better outcomes for ALL who are affected. Therefore, correcting nasal airflow often requires much more than adenoidectomy²⁵, however, this fact was lost for years among medical specialists. In the end, the results from T&A were limited, yet helped expose the fact that physicians did not yet have all the pieces to this puzzle, and we did not fully appreciate the complexity of the problem.

Shortly after all of this was happening in the dental world, we began acquiring significant experience in improving the nasal airway in adults via a minimally invasive surgical procedure called MIST (minimally invasive sinus technique) 26 . MIST was originally introduced in the mid-1990's by my mentor, Dr. Reuben Setliff. Reuben did not practice in an academic institution, had no fancy titles, yet he had a unique insight into nasal pathophysiology and how best to correct those suffering from all forms of inflammatory sinus disease. He was a superb surgeon with exceptional skills, insight, and intellect. His minimally invasive concepts have revolutionized nasal surgery and proved that less is often more when it comes to surgery in His techniques the nose. and philosophies were validated in adults and children²⁷, and provided a safe and effective platform upon which to expand our knowledge. The MIST procedure for airway correction often takes less

than 1 hour to complete, and allows patients to return home 2 hours after surgery, and back to work or school in 1 day. There is no external evidence of the surgery, no nasal packing, splints, or sutures, and minimal patient discomfort.

Correction of Nasal Obstruction

Since 2002, we have continued to modify the MIST procedure based on the critical review of our outcomes and feedback from our patients. In the mid2000's we introduced a simplified inoffice procedure for the correction of nasal valve collapse²⁸. In 2015 we published the first article on safe and effective ablation of the septal swell body²⁹ (Fig 11), which has greatly improved nasal breathing in adults and children. We have learned the true path of nasal breathing within the nose from computational flow studies performed by our colleagues in Singapore³⁰ and China³¹ (Fig 12).

Fig 11. Coronal CT image showing septal swell body



CT Showing Septal Swell Bodies *



Fig 12. Computational flow image of normal nasal airflow

Based on these collective advances, we now have the first reported results demonstrating that our minimally invasive nasal surgery, used alone as the sole intervention, can cure OSA in 48.7% of adults, regardless of the severity of their apnea, BMI, and gender³². An additional 43% of patients showed improvement in their AHI although not enough to be labeled a "cure".

Previously, nasal surgery alone had been shown ineffective in correcting OSA

with cure rates < 17%, and was primarily used to improve one's tolerance of CPAP³³. This has led sleep surgeons to pursue more aggressive and morbid procedures such as palatoplasty (UPPP), jaw advancement surgery, hyoid suspension, and robotic excision of the base of the tongue (known as TORS). These options are far from the minimally invasive goals of contemporary surgical practice, and proving unnecessary. Even the hypoglossal nerve stimulator. an expensive implant designed to stimulate

selective portions of the hypoglossal nerve during sleep to prevent hypopharyngeal obstruction, has success rates approaching $50\%^{34}$. The latter surgery takes about 3-5 hours to complete, prevents the use of MR imaging in the future, and is far more morbid and costly than MIST. Almost all the patients who were not cured after MIST had absolute or relative macroglossia, due to either a large tongue or small oral cavity. Our solution to tongue base prolapse is a device called Linguaflex. Linguaflex is an elastic tongue implant that is easily placed into the tongue in about 3-5 minutes using a straight needle 35 . Patients are unaware it is present once placed and it does not interfere with their speech or swallowing. It is adjustable, and easily reversible should the patient want it removed. This device is currently before the FDA awaiting 510K approval, and our data from clinical trials done in SE Asia show OSA cure rates from Linguaflex alone of over 70%. Combined with MIST, we believe we will have the ultimate minimally invasive options to cure OSA, SDB, and snoring in the majority of adult patients.

New Data Supports The Concept

Since 2015, we have applied these minimally invasive surgical principles to children with OSA and SDB, and continue to receive tremendous feedback from the children and their parents. With the use of a 3-dimensional CT imaging program (Fig 13), we have begun to measure the actual airway changes responsible for the subjective improvement in these children. We have found that both minimal throat area (2 dimensional measurement) and volume (3 dimensional space) have both increased an average of 42%, with *every* child studied showing improvement³⁶. We have also just completed another study measuring the volume changes in the nasal airway that occur after our modified MIST procedure, and the average child saw a 46% improvement nasal airway volume³⁷, which in translates into a 3-4 fold increase in nasal airflow! These findings were predictable based on the Starling Resistor Model of airway collapse and also validate the concept.



Fig 13. 3-D CT images can measure pharyngeal airway

(Fig 14) This model states that when one reduces nasal/upstream airway you will simultaneously resistance, reduce collapse of the pharyngeal airway by reducing negative intraluminal pressure. These ground-breaking results demonstrate that we can now accurately diagnose children with SDB, identify the cause of their compromised nasal breathing, appreciate the breathe and variation in their symptoms, and effective implement an treatment strategy that combines palatal expansion with minimally invasive nasal surgery, and T&A in selective individuals. Linguaflex will be a welcome addition to the adult population.

The last item to mention is VMR, which heretofore has been almost impossible to correct medically or surgically. However, in mid-2017, a company called Arrinex developed and received FDA approval for a new device called Clarifix³⁸. Using a cryo-probe strategically placed into the nose, the nerves responsible for autonomic control of nasal blood flow and secretions are literally frozen trans-mucosally in a 30

second application per side. The procedure can be performed in the office using local anesthesia in adults, and in the OR in children. Results are typically seen within 3-6 weeks and 2 year follow-up data shows over 90% success rates. Combined with MIST, we can now treat all causes of nasal obstruction in adults and children in a minimally invasive and effective manner.

Fig 14. Starling Resistor Model of Airway collapse

Starling Resistor Model of Airway Collapse



Eur Arch Otorhinolaryngol 2011:268;1365-73.

More Than Hope

This "perfect storm" of understanding the science, appreciating patient variability, creating adopting and technical innovations, and fostering learning peer-to-peer between disciplines, has given us the unique ability to confront and treat the pandemic that is OSA and SDB, and improve the quality of life for most who

suffer from them. Our work is just beginning, but we are confident we have "broken the code" on a condition that afflicts millions of people world wide, and is responsible for a huge health care burden for society. Today there is more than hope. References

- Jankowski, R., Nguyen, D. T., Poussel, M., Chenuel, B., Gallet, P., & Rumeau, C. (2016). Sinusology. European annals of otorhinolaryngology, head and neck diseases, 133(4), 263-268.
- Torre, C., & Guilleminault, C. (2018). Establishment of nasal breathing should be the ultimate goal to secure adequate craniofacial and airway development in children. Jornal de Pediatria (Versão em Português), 94(2), 101-103.
- Tay, D. K. L., & Pang, K. P. (2018). Clinical phenotype of South–East Asian temporomandibular disorder patients with upper airway resistance syndrome. Journal of oral rehabilitation, 45(1), 25-33.
- 4. Bharadwaj, R., Ravikumar, A., & N. Krishnaswamy, R. (2011). of Evaluation craniofacial morphology in patients with obstructive sleep apnea using lateral cephalometry and dynamic MRI. Indian Journal of Dental Research, 22(6), 739.

- Tolaymat, A., & Liu, Z. (2017). Sleep Disorders in Childhood Neurological Diseases. Children, 4(10), 84.
- Wexler, D., Braverman, I., & Amar, M. (2006). Histology of the nasal septal swell body (septal turbinate). Otolaryngology—Head and Neck Surgery, 134(4), 596-600.
- Ciolek, P. J., Xu, A., Anne, S., & Geelan-Hansen, K. (2017). Role of adenoidectomy in chronic nasal obstruction after nasal steroid therapy failure. American journal of otolaryngology, 38(3), 305-308.
- Davcheva-Chakar, M., Kaftandzhieva, A., & Zafirovska, B. (2015). Adenoid Vegetations– Reservoir of Bacteria for Chronic Otitis Media with Effusion and Chronic Rhinosinusitis. prilozi, 36(3), 71-76.
- Campos-Juanatey, F., Fernandez-Barriales, M., Gonzalez, M., & Portillo-Martin, J. A. (2017). Effects of obstructive sleep apnea and its treatment over the erectile function: a systematic review. Asian journal of andrology, 19(3), 303.
- 10. Hvolby, A. (2015). Associations of sleep disturbance with ADHD:

implications for treatment. ADHD Attention Deficit and Hyperactivity Disorders, 7(1), 1-18.

- Hawkins, S. M., Jensen, E. L., Simon,
 S. L., & Friedman, N. R. (2016).
 Correlates of pediatric CPAP adherence. Journal of clinical sleep medicine: JCSM: official publication of the American Academy of Sleep Medicine, 12(6), 879.
- 12. Muñoz, I. C. L., & Orta, P. B. (2014). Comparison of cephalometric patterns breathing in mouth and nose breathing children. International of journal pediatric otorhinolaryngology, 78(7), 1167-1172.
- 13. Šidlauskienė, М., Smailienė, D., Lopatienė, K., Čekanauskas, Е., Pribuišienė, R., & Šidlauskas, M. (2015). **Relationships** between malocclusion, body posture, and nasopharyngeal pathology in preorthodontic children. Medical science monitor: international medical journal of experimental and clinical research, 21, 1765.
- 14. Harvold, E. P., Vargervik, K., & Chierici, G. (1973). Primate experiments on oral sensation and

dental malocclusions. American journal of orthodontics, 63(5), 494-508.

- Ant, A., Kemaloglu, Y. K., Yilmaz, M., & Dilci, A. (2017). Craniofacial Deviations in the Children With Nasal Obstruction. Journal of Craniofacial Surgery, 28(3), 625-628.
- Shah, A., Shah, P., Goje, S. K., Shah, R., & Modi, B. (2017). Palatal Expansion and its Effects in Orthodontics. Advanced Journal of Graduate Research, 2(1), 31-36.
- Motro, M., Schauseil, M., Ludwig, B., Zorkun, B., Mainusch, S., Ateş, M., ... & Korbmacher-Steiner, H. (2016).
 Rapid-maxillary-expansion induced rhinological effects: a retrospective multicenter study. European Archives of Oto-Rhino-Laryngology, 273(3), 679-687.
- Bellerive, A., Montpetit, A., El-Khatib, H., Carra, M. C., Remise, C., Desplats, E., & Huynh, N. (2015). The effect of rapid palatal expansion on sleep bruxism in children. Sleep and Breathing, 19(4), 1265-1271.
- Goyal, A., Pakhare, A. P., Bhatt, G. C., Choudhary, B., & Patil, R. (2018). Association of pediatric obstructive

sleep apnea with poor academic performance: A school-based study from India. Lung India: Official Organ of Indian Chest Society, 35(2), 132.

- Owens, J. A. (2009). Neurocognitive and behavioral impact of sleep disordered breathing in children. Pediatric pulmonology, 44(5), 417-422.
- 21. Smith, D. L., Gozal, D., Hunter, S. J., & Kheirandish-Gozal, L. (2017).
 Parent-reported behavioral and psychiatric problems mediate the relationship between sleep-disordered breathing and cognitive deficits in school-aged children. Frontiers in neurology, 8, 410.
- 22. Zaffanello, M., Piacentini, G., Lippi, G., Fanos, V., Gasperi, E., & Nosetti, L. (2017). Obstructive sleepdisordered breathing, enuresis and combined disorders in children: chance or related association. Swiss Med Wkly, 147, w14400.
- 23. Laurikainen, E., Aitasalo, K., Erkinjuntti, M., & Wanne, O. (1992).
 Sleep apnea syndrome in children secondary to adenotonsillar

hypertrophy?. Acta Oto-Laryngologica, 112(sup492), 38-41.

- 24. Huang, Y. S., Guilleminault, C., Lee, L. A., Lin, C. H., & Hwang, F. M. (2014). Treatment outcomes of adenotonsillectomy for children with obstructive sleep apnea: a prospective longitudinal study. Sleep, 37(1), 71-76.
- Thadikonda, K. M., Shaffer, A. D., & Stapleton, A. L. (2018). Outcomes of adenoidectomy-alone in patients less than 3-years old. International Journal of Pediatric Otorhinolaryngology.
- 26. Catalano, P. J., Gupta, R. C., Warman, M., & Wijewickrama, R. C. (2014). Sinus Surgical Techniques from Caldwell-Luc to MIST. In Diseases of the Sinuses (pp. 389-410). Springer, New York, NY.
- 27. Setliff III, R. C. (1996). MIN IMALLY IN VASIVE SIN US SURGERY. Otolaryngologic Clinics of North America, 29(1), 115.
- Dolan, R. W., Catalano, P. J., Innis, W., & Wanees, E. (2009). In-office surgical repair of nasal valve stenosis. American journal of rhinology & allergy, 23(1), 111-114.

- Catalano, P., Ashmead, M. G., & Carlson, D. (2015). Radiofrequency ablation of septal swell body. Ann Otolaryngol Rhinol, 2(11), 1069.
- 30. De Yun Wang, H. P. L., & Gordon,
 B. R. (2012). Impacts of fluid dynamics simulation in study of nasal airflow physiology and pathophysiology in realistic human three-dimensional nose models. Clinical and experimental otorhinolaryngology, 5(4), 181.
- Tan, J., Han, D., Wang, J., Liu, T., Wang, T., Zang, H., ... & Wang, X. (2012). Numerical simulation of normal nasal cavity airflow in Chinese adult: a computational flow dynamics model. European archives of oto-rhino-laryngology, 269(3), 881-889.
- Peter Catalano Curative role of MIST in OSA (Publishing in progress).
- 33. Poirier, J., George, C., & Rotenberg,
 B. (2014). The effect of nasal surgery on nasal continuous positive airway pressure compliance. The Laryngoscope, 124(1), 317-319.

- Woodson, B. T., Strohl, K. P., Soose,
 R. J., Gillespie, M. B., Maurer, J. T.,
 de Vries, N., ... & Mickelson, S.
 (2018). Upper Airway Stimulation for
 Obstructive Sleep Apnea: 5-Year
 Outcomes. Otolaryngology–Head and
 Neck Surgery, 0194599818762383.
- 35. Catalano, P. J. (2014). U.S. Patent Application No. 14/055,159.
- 36. Alsufyani, N. A., Noga, M. L., Witmans, M., Cheng, I., El-Hakim, H., & Major, P. W. (2017). Using cone beam CT to assess the upper airway after surgery in children with sleep disordered breathing symptoms and maxillary-mandibular disproportions: a clinical pilot. Journal of Otolaryngology-Head & Neck Surgery, 46(1), 31.
- 37. Peter Catalano, Michael schlewet Nasal airway changes after nasal and sinus surgery in children with sleep disordered breathing (Publishing in progress).
- ClinicalTrials.gov Identifier: NCT03181594.