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A CRITICAL VIEW OF TONSILLITIS IN CHILDREN WITH IT'S DIAGNOSIS & TREATMENT APPROACH

Dr. Ritu Jyani*¹, Dr. Mohar² and Dr. Keerti Verma³

^{1,2}Second Year Post Graduate Scholar, Department of Kaumarbhritya, Rishikul Campus Uttarakhand Ayurved University, Haridwar, Uttarakhand, India.

³Head of Department of Kaumarbhritya Rishikul Campus Uttarakhand Ayurved University, Haridwar, Uttarakhand, India.

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*Corresponding Author Dr. Ritu Jyani

Second Year Post Graduate Scholar, Department of Kaumarbhritya, Rishikul Campus Uttarakhand Ayurved University, Haridwar, Uttarakhand, India.

ABSTRACT

Tonsillitis is one of the most widely conditions that affect children in all ages during the childhood. The overall management of tonsillitis is complicated related to it's causes, specific diagnosis and appropriate treatment measures. Therefore, the aim of this review was to highlight the tonsillitis in children with particular diagnostic approach and treatment measure that are currently worldwide used to deal with pediatric tonsillitis.

KEYWORDS: Tonsillitis; Tonsillectomy, Tonsils; Streptococci, Mononucleosis, Sore throat.

INTRODUCTION

Tonsillitis or throat infection is one of the most common health problems worldwide. [1-3] Tonsillitis is severe intermittent throat

infections, which is characterized by <5 episodes of tonsillitis in a year. Beside this acteria as the main causes of the tonsillitis, some viruses & infectious mononucleosis, can be possible causes. Diagnosis of tonsillitis is clinicaly/laboratory, though sometimes, it can be a challenging to differentiate between viral & bacterial infections. As more accurate tests take longer time to conclude the results, rapid antigen testing with a very low sensitivity is repeat in the diagnosis of bacterial tonsillitis. Other causes include infectious mononucleosis from Epstein Barrvirus (EBV) infection, cytomegalovirus(CMV), human immuno deficiency virus (HIV), hepatitis A, rubella and toxoplasmosis. [4] Tonsillectomy is still one of the atmost common surgical procedures for

the treatment of tonsillitis in children. Tonsillitis and tonsillectomy are likely to have a distinct link where the incidence of one should reflect that of the other. While the entirely incidence of tonsillectomy in a population may be extreme less than the total incidence of tonsillitis, a healthcare system should be capable to familiarize up to surge rates of a specific surgical treatment when the specific sign for that treatment up surges.^[5] However, the aim of this review to provide a valuable updates in the diagnosis and treatment approach in childhood tonsillitis.

Tonsils

The tonsils contain immune-cells such as B-lymphocytes, T-lymphocytes and other antigen cells such as macrophages, which serves as an immune acquisition and defense. [6-8] Therefore, the core phase of the immune acquisition carry on upto the age of six, the palatine tonsils are physiologically hyperplastic at this stage. [9] At that time there is a shrinkage in are gression up to the age of 12. [10,11] The palatine tonsils have a forceful blood flow from four diversent vessels. These vessels give out primarily to the superiorand inferior tonsil pole, as well as the particular center of the tonsils sideways. [6] The tonsils have profound crypts to retain the organ exterior as huge as possible and to generate a detaching surface for potential antigens. [12] The crypts and their neighboring germinal centers are disconnected from each other by suitable connective tissue. [13]

Etiology of Tonsillitis

The majority of cases of tonsillitis are caused by bacterial particularly beta-hemolytic and others treptococci. However, in tonsillitis related to infectiou smononucleosis, the most common virus is the EBV, which present in 50% of children. CMV, hepatitis A, HIV, rubella and to xoplasmosis infections may also result in the clinical picture of in factious mononucleosis, which requires differential diagnosis. [4] With regard to bacterial in fection, severalgerms were continuously in criminateddue continuous advancement in detection methods. [1416] At the time of discovery of rheumatic fever, all cases of upper respiratory tract infections were thought to becaused by group Astreptococcus [17,18], but later on there are anaerobes, suchas *Fusobacteriumn ecrophorum*, *Streptococcus intermedius* and *Prevotella melaninogenica* and *his ticola* were incriminated. [19,21] When investigate the organisms that in fect the tonsilin different phase soft tonsillar life. Accurately, itwas discovered that children before the age of eight years old have a tendency in the course of adiffuse, intracellular organisms augmentation with interstitial

abscesses^[22], While a superficial bacterial accumulation at the edge of the crypts was verified in adolescents.^[23] The oral cavity and specially the furrowed tonsil is a reservoir for multiple pathogens including viruses, bacteria, parasites^[24] and fungi.^[25] These microbes belong to the transient flora and human lives in cooperation with them.^[19,26] However, more than 100 bacteria can be detected in the tonsils of children and adults with and without recurrent tonsillitis. Moreover, about 52 different bacterial strains can be identified in each patient, whether child or adult, whether ill or healthy. These bacterial strains represent 90% of the to talpathogen load.^[19] In thea cute tonsillitis in children, the streptococcus was found to constitute up to 30% ^[27], followed by *Haemophilus influenzae* and *Neisseria*.^[19] However, mixedin fections (both bacterial and viral) may show similar clinical features.^[28]

Types of Tonsillitis

There are several type soft on sillitisacute, recurrent, and chronic tonsillitis, and peritonsillar abscess.

Acute Tonsillitis

It is also known as severe tonsillitis^[29] oracutesorethroat^[4], is inflammation of the tonsils, which is caused by bacteria or virus with odynophagia. The condition is characterized by swelling and redness of the tonsils, may be with exudate, cervical lymphadenopathy and fever>38.3°Crectal^[30,31]

The odynophagia for 24 to 48 hours, resembles the symptoms of a common cold is excepted from the definition of acute tonsillitis.^[4] Acute tonsillitis is an inflammatory process of the tonsillart issues and is generally infectious insource. It is part of the spectrum of pharyngitis, which varies from localized tonsils' infection to widespread infection of the pharynx and usually affects young healthy adults. Simple sore throats secondary to viral orbacterial pharyngitis are mutual and commonly do notneed hospital admission or antibiotic treatment. Caring management such as analgesia and sufficient hydration issusually necessary.^[32] It is essential to differentiate the catarrhal angina with redness and swelling of the tonsils(earlystage) from the follicul arangin a with stipple like fibrin deposits from the lacunarang in a with confluent eposits(latestage). This differentiation should be based on the stage and look of the deposits, or the exudate on the tonsils.^[33-38,31] Acute infections of the palatine tonsils mostly arise in children at schoolage, but patients of any age may be affected. Tonsillitis of viral infection is

regularly treated with supportive care. Bacteria ltonsillitisis most or dinarily caused by Strep to coccus pyogenes. Poly-microbial infections and viral pathogens are also significant causes of infection.^[39]

RecurringTonsillitis

Recurrenttonsillitisorrecurrentthroatin fections^[4], denotes recurrences of acute tonsillitis. This type of tonsillitisis defined ifepisodes of tonsillitis in one year evidenced in seven bacterial cultures, or five in two sequential years or three in three repeated years. Insuch conditions, the pause of the antibiotic leads to another bout of the bacterial in fection with in a few weeks, thus triggering it to return oncemore.^[40,41]

On the other hand a single violence of acutetonsillitiscanbec a used by several diverse bacterial organisms^[19,22] and flash up again a few weeks after termination of an antibiotic treatment.^[42] Relevant on the rate of recurrence and severity of such episodes, there isaclue for tonsillectomy.

Chronic Tonsillitis

In this type is associated with chronic sorethroat, in which the infection causes recurrent tonsillitis. This isusu all yassociated with bad breath type and persistenttendercervicalnodes. Chronictonsillitis describe the most common lesions contained by pharynxinflammatory pathology with multiple complications both local regional and atthed istance(glomerulonephritis, join trheumatism, endocarditis, enteritis, etc.). [43-46] Chronic tonsillitis can be also the site of some specific infections such as tuberculosis and syphilitic. [45,46] Chronictonsillit is usually describe focal tonsillitis, hypertrophic or scleroatrophiccaseous cryptic tonsillitis as recurrent type, and simple hypertrophic tonsillitis soft type in children and hardtypeinadults. [46] In most of the cases, the hypertrophic form occurs in adult and elder child, the tonsils were hypertrophied, congested, with reduced flexibility in the amygdalian space with apparent cryptseradicatingspontaneouscaseum, but also when they were compelled by spatula on the anterior pilier. [47]

Peritonsillar Abscess(PTA) or Quinsy

It is an acute tonsillitis with formation of anabscess, typically on one side.^[48] The intratonsillar, para-/peritonsillarorretrotonsillarspaces may be associated with abscess formation. When the acutetonsillitis is leftuntreated, the bacterial infection usually lead to

peritonsillarabscess, which develops lateral to the tonsillar region. The area of the peritoneal ciliates is clearly visible as a swollen area with pus accumulation. Staphylococci, Streptococci, Haemophilus and Fusobacterium necrophorumarethe most common pathogens responsible of peritonsillarabscess. Novirusisinvolved. Altered voice quality, uneaseofmouthopening, coarsebreath, fever and severe sore throat are the main symptoms. [49-

Diagnosis of Tonsillitis

The diagnosis softonsillitis in children and adults usually start with clinical diagnosis.^[51] Diagnosis of tonsillitis is based on history to find out whether tonsillitis is recurrent and a physical examination of the throat. This can give an clue whe ther the cause is bacterial or viral or other. [52] Pain, fever, primarily cough, hoarseness, and rhinorrhoea frequently occur inviral tonsillitis, while pain with lymph node swelling canoccurin bacterial tonsillitis with mainly tonsils' exudate and fever >38.3°C. [30] Although strep to coccalantigen test is lesssensitive, but it can confirm the diagnosis of strep to cocci with 98% specificity. However, it was recommended that negative tests should be further confirmed with other more specifictests. Of the drawback of bacterial and viral RNA rapidtests, they can be positive in 10% of healthy children(usually asymptomatic chronic carriers of staphylococci and streptococci), who are absolutely don't require treatment. [53-There fore, smears or these tests should only be performed on symptomatic patients. [56] At the initial phase, it is difficult to distinguish between viral and bacterial tonsillitis, particularly on considering that around 97.5% of cases, at least harbor one virus, eveninthebacterialtonsillitis, adenovirusandparainfluenza virus may be found in about 47.5% of thecases. [57-59] However, it is recommended to get a tonsillarswab for rapid antigen testing rapid antigen detection(RAD) in children or adolescents with a history, signsand/orsymptomsofsuspectedinfectionl or suspicion of infection, a bacterial cultures hould be done. In the case of appositive RAD test results, the bacterial culture is not compulsory for the high reliability and specificity of the tests. [60-62]

The dosage of the anti-strep to coccusantibodies Anti-streptolysinO(ASOT) is not recommended in the routine diagnosis of strep to coccalpharyng it is since the existence of the seantibodies reveals past infections and rather than on going infections. [63] Patients with raised ASO titers and recurrent tonsillitis episodes are known to be at greater risk for rheumatic heart disease. [64] Higher ASO titers can be detected in different clinical disorders other than the classic post-streptococcal related infections. In such ases it is not essential to be correlated with positive culture and or with in flammatory parameters. ^[65]

Furthermore, differential diagnosis sometimes deemed important. Tonsils hyperplasia, dry air flows associated with turbinate hyperplasia [66], and allergy associated with chronic inflammation^[67], in particulardust mite and mould allergy.^[68] Tooth decay can also lead to recurrent to nsillitis^[69] and it was found that tonsil removal can positive lyaffectpediatric periodontal disease.^[70]

Inunilateral tonsillar hyperplasia, the differential diagnosis of lymphoma should be considered. [71] In pediatric unilateral tonsillitis, the differential diagnosis of Plaut-Vincent angina (caused by the spirochete (Treponemavincentii and Fusobacteriumnucleatum)) shoul dberolledout.[72]

Inacuteton sillitis, the diagnosis of scarlet fever produced by streptococci group A, which is responsible for the productionerythrogenictoxin, which result stypical rashforscarlet fever. [73] Mononucleosis is caused by the EBV, particularly in children under the age of 10 years old. Although the disease is usually symptomless, but flu-like signs can occur in elderly people. Sometimes the disease occurs again course with severe sorethroatandswollen, coated tonsils and largelymphnodes, particularlyinadolescents. Moreover, splenic enlargement, loss of appetite, chills, dry cough, nausea and nights weatscanoccur. However, the disease can occur again. At present, there is still no approved vaccine for the EBV. [75,76]

Treatment of tonsillitis Antibiotic treatment

Penicillin still the treatment of choice for Streptococcuspyogenestonsillitis, and amplified aminoopenicillins have grown usefulness in performance with the growing incidence of betalactamase producingbacteria. [39]

As a beta-lactam antibiotic therapy offers quite reliable protection against the feared rheumatic fever and glomerulonephritis. These conditions often lead specifically in developing countries to arthritis, myocarditis and death. [77]

The penicillin, especially in children and doles cents, display the highest advantage for it being the lowest cost. Cephalosporin is, however, more effective in children under the age of 12 years and for chronic recurrent tonsillitis, as it can destroy more strains of streptococci. [78] Moreover, the short-term therapy with azithromycin(20mg/kg)[80] for three days or clarithromycin and cephalosporin for five days is equivalent to the long-term penicillin therapy with improved amenability.^[81]

Analgesics

For acute tonsillitis, the most common Non-steroidal anti-inflammatory is ibuprofen, which shows the top efficacy with the least side effects compared with paracetamol and acetylsalicylic acid. Extra benefit of ibuprofen is extended period of action of 6-8 hours in comparison to paracetamol. Diclofenac sodium and ketorolac in children have less cut off sites and are metabolized rapidly, which necessitate the adjustment of the dose (higher do sage than in adults). Metamizol should be avoid edasananalgesic in children because of the small but existing risk of agranulocytosis. [85]

Steroids

The use of steroids in children display sasubstantial enhancement in symptoms with slight side effects and without any effect son disease evolution. The best consequences were realized in verified streptococcalpharyngitis for dexamethasone as well as betamethasone and prednisolone with a perfect decrease in the pain and feeling of illness that associated with acute tonsillitis. [87]

Mouthwashes

Antiseptic mouthwashes with chlorhexidine or benzydamine show signs of relief in children and adults.^[88] Typical herbal gargles contain sage, thyme and chamomile, can lubricate and preserve the mucousmembranes. However, several substance scontaining ethanol as an extraction solvent and are not approved for children<12 years old. Nasturtium and horseradishroot are contained in some pharmaceutical drugs, which have antimicrobial, antiviral and antifungal properties.^[89]

Tonsillectomy

Is still one of the most common tonsillar surgery during childhood. A tonsillectomy in children before the age 6 years old should only be performed if the child suffers from recurrent acute bacterial tonsillitis. In all other cases such as hyperplasia of the tonsils, the low risk partial onsillecto my should be the first line treatment. Postoperative pain and the risk of hemorrhage are lesserinpartial tonsillectomy. Regardless of the fact that, tonsillotomy is performed by laser, radio frequency, shaver, coblation, bipolar scissor or Colorado needle, open crypt and tonsil tissue that is left behind should be considered. Total extra capsular

tonsillectomy is still considered in severely affected children with recurrent infections of the tonsils, allergy to antibiotics, PFAPA syndrome(periodic fever, aphthousstomatitis, pharyngitis, and cervicaladenitis) and peritonsillarabscess. With regard to the frequency and seriousness of the recurrent tonsillitis the sign for tonsillectomy in children is necessary if seven or more well documented, clinically significant, sufficiently treated episodes of throat infection happen in the past year, or five or more such episodes happen in each of the two previous years. Post-tonsillectomy bleeding may continue till the entire wound is totally healed, which is usually within three weeks. Life-threatening hemorrhages may increase after minor bleedings, which can spontaneously stop in most of the cases of atal outcome after tonsillectomy, due to correct management of hemorrhage. In more younger children hemorrhage can be life threatening due to the lower blood volume and the danger of aspiration with asphyxia. All "hot" procedures with laser, radiofrequency, coblation, monoorbipolar forceps have a greater risk of late hemorrhage. The preoperative information about the surgery should be bedone with the child and the parents in a quiet and objective atmosphere with a written consent. [77] Intracapsular/subcapsularorsubtotalton sillecto my:involves the removal of the lymphatic active tissue of the tonsil, including all crypts and follicles. However, the intracapsular tonsillectomy is partially associated with tonsillotomy. [90]

CONCLUSION

Bacteria is still the most causative agent with widely used antigen tests for confirmation. Tonsillectomy is still one of the most frequently performed surgical interventions in children.

REFERENCES

- 1. Lau, A. S., Upile, N. S., Wilkie, M. D., Leong, S.C., & Swift, A.C. Ther is ingrate of admissions for tonsillitis and neck space abscesses in England, 1991-2011. The Annals of The Royal College of Surgeons of England, 2014; 96(4): 307-310.
- 2. Klug, T. E. Peritonsillar abscess: clinical aspects of microbiology, riskfactors, and the association with parapharyngeal abs cess. Clin Infect Dis, 2009; 49: 1467-72.
- 3. Borgström, A., Nerfeldt, P., Friberg, D., Sunnergren, O., & Stalfors, J. Trends and changes in paediatric tonsil surgery in Sweden 1987–2013: a population-based cohort study. BMJopen, 2017; 7(1): e013346.
- 4. Georgalas, C.C., Tolley, N.S., & Narula, A .Searchdate March 2009.

- 5. Millington, A. J., & Phillips, J. S. Current trends in tonsillitis and tonsillectomy. *The Annals of The Royal College of Surgeons of England*, 2014; *96*(8): 586-589.
- 6. Westermann, J. Organedes Abwehrsystems. In *Anatomie* (355-375). Springer Berlin Heidelberg, 2010.
- 7. Nave, H., Gebert, A., & Pabst, R.(2001). Morphology and immunology of the humanpalatinetonsil. *Anatomyandembryology*, 204(5): 367-373.
- 8. Brandtzaeg, P.(2011). Immune function sofnasopharyngeallymphoidt issue. In *RecentAdvances in Tonsils and Mucosal Barriers of the UpperAirways* (Vol.72, pp.20-24). KargerPublishers.
- Kaygusuz, I., Gödekmerdan, A., Karlidağ, T., Keleş, E., Yalçin, Ş., Aral, I., & Yildiz, M. (2003). Early stage impacts of tonsillectomy on immunefunctionsofchildren.
 International journal of pediatricotorhinolaryngology, 67(12): 1311-1315.
- 10. Kaygusuz, I., Alpay, H.C., Gödekmerdan, A., Karlidag, T., Keles, E., Yalcin, S., & Demir, N.(2009). Evaluation of long-termimpacts of ton sillectomy on immune functions of children: afollow-up study. *International journal of pediatricotorhinolaryngology*, 73(3): 445-449.
- 11. Perry, M. E., & Slipka, J. (1993). Formation of thetonsillar corpuscle. *Functional and developmentalmorphology*, *3*(3): 165-168.
- 12. Drucker, M., Drucker, I., Neter, E., Bernstien, J., & Ogra, P.L.(1978). Cellmediatedimmuneresponses to bacterial antigens on human mucosalsurfaces. In *Secretory Immunity and Infection* (pp.479-488). Springer, Boston, MA.
- 13. Perry, M. E., Jones, M. M., & Mustafa, Y. (1988). Structure of the crypt epithelium in human palatinetonsils. *Acta Oto-Laryngologica*, *105*(sup 454): 53-59.
- 14. Al-Mazrou, K.A., & Al-Khattaf, A.S. (2008). Adherentbiofilmsinadenotonsillar diseases in children. *Archivesof Otolaryngology–Head & Neck Surgery*, 134(1): 20-23.
- 15. Drago, L., De Vecchi, E., Torretta, S., Mattina, R., Marchisio, P., & Pignataro, L.(2012). Biofilmformation by bacteria isolated from upper respiratory tract before and after adenoton sillectomy. *Apmis*, *120*(5): 410-416.
- 16. Jensen, A., Fagö-Olsen, H., Sørensen, C.H., & Kilian, M. (2013). Molecular mapping to specieslevel of the tonsillar crypt microbiota associated with health and recurrent tonsillitis. *PLoSOne*, 8(2): e56418.
- 17. MatanoskiGM, PriceWH, FerenczC.(1968). Epidemiologyofstreptococcalinfectionsinrheumaticandnon-rheumaticfamilies.II.

- Theinter-relationship of streptococcal infections to age, family transmission and type of group A. Am JEpidemiol, 87(1): 190-206.
- 18. FeinsteinAR, LevittM.(1970). Tonsilsandrheumaticfever. N Engl J Med., 2; 282(14): 814.
- 19. Jensen, A., Fagö-Olsen, H., Sørensen, C.H., & Kilian, M. (2013). Molecular mapping to species level of the tonsillar crypt microbiota associated with health and recurrent tonsillitis. PLoSOne, 8(2): e56418.
- T.E., K., 20. Klug, Henriksen, J.J., Fuursted, & Ovesen, T.(2011).Significantpathogensinperitonsillar abscesses. European journal of clinical microbiology & infectious diseases, 30(5): 619-627.
- 21. Klug, T.E., Henriksen, J.J., Fuursted, K., & Ovesen, T.(2011). Similarrecoveryrates of Fusobacteriumnecrophorumfromrecurrentlyinfected and non-infected tonsils. Dan MedBull, 58(7): A4295.
- 22. Swidsinski, A., Göktas, Ö., Bessler, C., Loening-Baucke, V., Hale, L. P., Andree, H., ... & Lochs, H.(2007). Spatialorganisationofmicrobiotainquiescentadenoiditis and tonsillitis. Journal of clinical pathology, 60(3): 253-260.
- 23. Ramírez-Camacho, R., González-Tallón, A.I., Gómez, D., Trinidad, A., Ibáñez, A., García-Berrocal, J.R.,... & SanRomán, J.(2008). Environmentalscanningelectronmicros copy for biofilm detection in tonsils. Acta Otorrinolaringologica (English Edition), 59(1): 16-20.
- W., C., 24. Mücke, Huber, H. & U. The Ritter, (1994).microbecolonizationofthepalatinetonsils of healthy school children. age Zentralblattfur Hygieneund Umweltmedizin=International journal of hygiene and environmental medicine, 196(1): 70-74.
- I.J., Reilly, P.G., Hay, A.J., & 25. Mitchelmore, Tabaqchali, S.(1994). Tonsilsurfaceandcoreculturesinrecurrenttonsillitis: prevalenceofanaerobesandbetalactamaseproducingorganisms. European Journal of Clinical Microbiology & *Infectious Diseases*, *13*(7): 542-548.
- 26. Schwaab, M., Gurr, A., Hansen, S., Minovi, A. M., Thomas, J. P., Sudhoff, H., & S. (2010). Humanβ-Defensinsindifferent states of diseases of the tonsillapalatina. European Archives of Oto-Rhino-Laryngology, 267(5): 821-830.
- 27. Brook, I., & Foote, P. A. (1986). Comparison of the microbiology of recurrenttonsillitis between children and adults. The Laryngoscope, 96(12): 1385-1388.

- 28. Proenca-Modena, J. L., Valera, F. C. P., Jacob, M.G., Buzatto, G. P., Saturno, T. H., Lopes, L., ... & Tamashiro, E. (2012). High rates of detection of respiratory virusesintonsillart issues from children with chronic adenotonsillar disease. *PLoSOne*, 7(8): e42136.
- 29. Lange, G. (1973). Angina and chronic tonsillitis--indicationsfortonsillectomy. *Zeitschriftfür Allgemeinmedizin*, 49(8): 366.
- 30. Stelter, K.(2014). Erkrankungender Gaumenmandelnim Kindesalter. *Laryngo-rhino-otologie*, *93*(S01): S84-S102.
- 31. Stuck, B.A., Windfuhr, J.P., Genzwürker, H., Schroten, H., Tenenbaum, T., & Götte, K. (2008). Tonsillectomy in children. *Deutsches Ärzteblatt International*, 105(49): 852.
- 32. Bartlett, A., Bola, S., & Williams, R. (2015). Acutetonsillitis and its complications: an overview. *Journal of the Royal Naval Medical Service*, *101*(1): 69-73.
- 33. Berghaus, A. (2005). Chronic inflammation of the upper airways. Operation instead of antibiotic. *MMW Fortschritte derMedizin*, 147(39): 27.
- 34. Berghaus A, Pirsig W. (1996). Mundhöhleund Pharynx. In: Berghaus A, Rettinger G, Böhme G, editors. Hals-Nasen-Ohren-Heilkunde. 1st ed. Stuttgart: Hippokrates Verlag;
- 35. Borschmann, M. E., & Berkowitz, R. G. (2006). One-offstreptococcalserologictesting in young children with recurrenttonsillitis. *Annals of Otology, Rhinology & Laryngology*, 115(5): 357-360.
- 36. Fujikawa, S., Hanawa, Y., Ito, H., Ohkuni, M., Todome, Y., & Ohkuni, H. (1988). Streptococcalantibody: asanindicatoroftonsillectomy. *ActaOto-Laryngologica*, *105*(sup454): 286-291.
- 37. Kurien, M., Stanis, A., Job, A., & Thomas, K.(2000). Throat swab in the chronic tonsillitis: howreliableandvalidisit?. *Singaporemedicaljournal*, 41(7): 324-326.
- 38. Stuck BA, Genzwürker HV. (2008) Tonsillectomyin children: preoperative evaluation of risk factors. Anaesthesist, 57(5): 499-504.
- 39. SidellD, ShapiroNL.(2012). Acutetonsillitis. InfectDisord DrugTargets, 12(4): 271-6.
- 40. Szpunar, J., & RYBAKOWA, M.(1961). Electrophoretic Serum Studies Studies of Childrenwith Frequently Recurring AcuteTonsillitis. *Archives of Otolaryngology*, 74(3): 267-271.
- 41. Senska, G., Ellermann, S., Ernst, S., Lax, H., & Dost, P.(2010). Recurrenttonsillitisinadults: qualityoflifeaftertonsillectomy. *Deutsches Ärzteblatt International*, 107(36): 622.

- 42. JENSEN, J.H., & LARSEN, S.B.(1991). Treatment of recurrentacutetonsillitis with clindamycin. An alternative to tonsillectomy?. *Clinical Otolaryngology*, *16*(5): 498-500.
- 43. Gaffney, R.J., & Cafferkey, M.T.(1998). Bacteriology of normal and disease dtonsilsassessedbyfine-needleaspiration: Haemophilusinfluenzae and the pathogenesis of recurrentacute tonsillitis. *Clinicalotolaryngology and allied sciences*, 23(2): 181-185.
- 44. Mattila, P.S., Tahkokallio, O., Tarkkanen, J., Pitkäniemi, J., Karvonen, M., & Tuomilehto, J.(2001). Causes of tonsillar disease and frequencyoftonsillectomyoperations. *Archives of Otolaryngology–Head & Neck Surgery*, 127(1): 37-44.
- 45. Bailey, B.J., Johnson, J.T., Newlands, S.D., Calhoun, K. H., Curtin, H. D., & Deskin, R.W.(2006). Diagnosis and treatment of thyroidandpara thyroid disorders. *Head & Neck Surgery Otolaryngology. Fourthedition. Philadelphia: Lippincott Williams & Wilkins*, 1630.
- 46. Bourbeau, P. P. (2003). Role of the microbiology laboratory in diagnosis and management of pharyngitis. *Journal of clinicalmicrobiology*, 41(8): 3467-3472.
- 47. Mogoantă, C. A., Ioniță, E., Pirici, D., Mitroi, M., Anghelina, F., Ciolofan, S., & Pătru, E.M.I.L.I. (2008). Chronictonsillitis: histologicalandimmunohistochemicalaspects. *Rom J Morphol Embryol*, 49(3): 381-386.
- 48. Dalton, R. E., Abedi, E., & Sismanis, A. (1985). Bilateralperitonsillarabscesses and quinsy tonsillectomy. *Journal of the National Medical Association*, 77(10): 807.
- 49. Mazur, E., Czerwińska, E., Korona-Głowniak, I., Grochowalska, A., & Kozioł-Montewka, M.(2015). Epidemiology, clinicalhistoryandmicrobiologyofperitonsillarabscess. European Journal of Clinical Microbiology & Infectious Diseases, 34(3): 549-554.
- 50. WangY-P, WangM-C, LinH-C, ChouP. The Impact of Prior Tonsillitis and Treatment Modalityonthe Recurrence of Peritonsillar Abscess: A Nationwide Cohort Study. AmarS, ed. *PLoSONE*. 2014; 9(10): e109887.doi:10.1371/journal.pone.0109887.
- 51. Klug, T. E. (2009). Peritonsillar abscess: clinical aspects of microbiology, risk factors, and the association with parapharyngealabscess. *ClinInfectDis*, 49: 1467-72.
- 52. Proenca-Modena, J. L., Valera, F. C. P., Jacob, M.G., Buzatto, G. P., Saturno, T. H.,

- Lopes, L., ... & Tamashiro, E. (2012). High rates of detection of respiratory viruses in tonsillar tissues from children with chronic adenotonsillar disease. *PLoSOne*, 7(8): e42136.
- 53. Tanz, R. R., & Shulman, S.T. (2007). Chronicpharyngeal carriage of group A streptococci. *The Pediatric infectious disease journal*, 26(2): 175-176.
- 54. WesselsMR.(2011). Clinical practice. Streptococcal pharyngitis. NEnglJMed, 17; 364(7): 648-55.
- 55. Roberts, A.L., Connolly, K.L., Kirse, D.J., Evans, K., Poehling, K. A., Peters, T. R., & Reid, S.D. (2012). Detection of group A Streptococcusintonsils from pediatric patients reveals high rate of asymptomatic streptococcal carriage. *BMC pediatrics*, 12(1): 3.
- 56. Shulman, S.T., Bisno, A.L., Clegg, H.W., Gerber, M.A., Kaplan, E.L., Lee, G.,...& VanBeneden, (2012). Clinical practice guideline for the diagnosis and management of group Astreptococcalpharyngitis: 2012 update by the Infectious Diseases Society of America. *Clinicalinfectious diseases*, 55(10): e86-e102.
- 57. Drago, L., Esposito, S., De Vecchi, E., Marchisio, P., Blasi, F., Baggi, E., ... & Pignataro, L. (2008). Detectionofrespiratoryvirusesandatypicalbacteria in children's tonsils and adenoids. *Journal of clinical microbiology*, 46(1): 369-370.
- 58. Piacentini, G.L., Peroni, D.G., Blasi, F., Pescollderungg, L., Goller, P., Gallmetzer, L., ... & Boner, A. L. (2010). Atypical bacteria in adenoids and tonsils of children requiringadenotonsillectomy. *Acta oto-laryngologica*, *130*(5): 620-625.
- 59. Proenca-Modena, J. L., Valera, F. C. P., Jacob, M.G., Buzatto, G. P., Saturno, T. H., Lopes, L., ... & Tamashiro, E. (2012). High rates of detection of respiratory viruses in tonsillar tissues from children with chronic adenotonsillar disease. *PLoSOne*, 7(8): e42136.
- 60. Gurol, Y., Akan, H., Izbİrak, G., Tekkanat, Z. T., Gunduz, T. S., Hayran, O., & Yilmaz, G. (2010). The sensitivity and the specifity of rapid antigentestinstreptococcalupperrespiratorytract infections. *International journal of pediatricotorhinolaryngology*, 74(6): 591-593.
- 61. Pelucchi, C., Grigoryan, L., Galeone, C., Esposito, S., Huovinen, P., Little, P., & Verheij, T. (2012). Guideline for the management of acute sore throat. *Clinical microbiology and infection*, 18(s1): 1-28.
- 62. Tajbakhsh, S., Gharibi, S., Zandi, K., Yaghobi, R., & Asayesh, G.(2011). Rapiddetection of Streptococcus pyogenes in throat swab specimens by

- fluorescentinsituhybridization. *European review for medical and pharmacological sciences*, 15(3): 313-317.
- 63. Chiappini, E., Principi, N., Mansi, N., Serra, A., DeMasi, S., Camaioni, A.,... & Speciale, A.M.(2012). Management of acutepharyngitisin children: summary of the Italian National Institute of Health guidelines. *Clinicaltherapeutics*, 34(6): 1442-1458.
- 64. Sahin, M. S., Yalcin, M. U., &Kocyigit, D. (2016). Prevalence of rheumatic heart disease in patientswithrecurrenttonsillitisandelevatedanti-streptolysinOtiters. *International journal of pediatricotorhinolaryngology*, 89: 133-135.
- 65. Ben-Chetrit, E., Moses, A. E., Agmon-Levin, N., Block, C., & Ben-Chetrit, E. (2012). Serum Levels Ofanti-streptolysin Oantibodies: the irroleinevaluatingrheumatic diseases. *International journal of rheumatic diseases*, 15(1): 78-85.
- 66. Berkowitz, R.G., & Mahadevan, M.(1999). Unilateraltonsillarenlargementandtonsillarlymphomain children. *Annals of Otology, Rhinology & Laryngology*, 108(9): 876-879.
- 67. Sadeghi-Shabestari, M., Moghaddam, Y.J., & Ghaharri, H.(2011). Is there any correlation between allergy and adenotonsillartissuehypertrophy?. *International journal of pediatricotorhinolaryngology*, 75(4): 589-591.
- 68. Karaca, Ç. T., Toros, S. Z., Noseri, H., Külekçi, S., Kalayck, Ç., Oysu, Ç., ... & Egeli, E. (2012). Roleofallergyin children with adenotonsillarhypertrophy. *Journal of Craniofacial Surgery*, 23(6): e611-e613.
- 69. Georgalas, C., Kanagalingam, J., Zainal, A., Ahmed, H., Singh, A., & Patel, K. S. (2002). The association between periodontal disease and peritonsillar infection: a prospective study. *Otolaryngology—Head and NeckSurgery*, 126(1): 91-94.
- 70. Demir, U.L., Cetinkaya, B., Karaca, S., & Sigirli, (2013). The impacts of adenotonsillarhypertrophy on periodontal health in children: aprospective controlled pilot study. *American journal of otolaryngology*, *34*(5): 501-504.
- 71. Berkowitz, R.G., & Mahadevan, M.(1999). Unilateraltonsillarenlargementandtonsillarlymphomain children. *Annals of Otology, Rhinology & Laryngology*, 108(9): 876-879.
- 72. Gebhardt, B., Herrmann, K., Roessner, A., & Vorwerk, U.(2010). Differenzialdiagnostikdernekrotisierenden Tonsillitis. *Laryngo-rhino-otologie*, 89(05): 266-269.

- 73. Lee, P. K., & Schlievert, P. M. (1991). Moleculargenetics of pyrogenic exotoxin "superantigens" of group A streptococci and Staphylococcus aureus. In *Superantigens*(pp.1-19). SpringerBerlinHeidelberg.
- 74. Walther, L. E., Ilgner, J., Oehme, A., Schmidt, P., Sellhaus, B., Gudziol, H.,... & Westhofen, M.(2005). Die infektiöseMononukleose. *Hno*, 53(4): 383-394.
- 75. Cohen, J.I., Mocarski, E.S., Raab-Traub, N., Corey, L., & Nabel, G. J. (2013). The need and challenges for developmentofan Epstein-Barrvirusvaccine. *Vaccine*, *31*: B194-B196.
- Ruiss, R., Jochum, S., Wanner, G., Reisbach, G., Hammerschmidt, W., & Zeidler, R.(2011). Avirus-likeparticle-basedEpstein-Barrvirusvaccine. *Journal of virology*, 85(24): 13105-13113.
- 77. Stelter, K.(2014). Tonsillitisandsorethroatinc Tonsillitis and sorethroatin children. *GMS current topics inotorhinolaryngology, headandnecksurgery, 13*.
- 78. Casey, J. R., & Pichichero, M. E. (2004). Meta-analysis of cephalosporinversuspenicillin treatment of group Astreptococcaltonsillopharyngitisin children. *Pediatrics*, 113(4): 866-882.
- 79. Chiappini, E., Principi, N., Mansi, N., Serra, A., DeMasi, S., Camaioni, A.,... & Speciale, A.M. (2Cohen, R., Reinert, P., De La Rocque, F., Levy, C., Boucherat, M., Robert, M., ... & Bingen, E. (2002). Comparison of two dosages of azithromycin for three penicillin V for in days versus ten days acute group Astreptococcaltonsillopharyngitis. The Pediatricinfectious disease journal, 21(4): 297-303.
- 80. Altamimi, S., Khalil, A., Khalaiwi, K.A., Milner, R. A., Pusic, M. V., & Al Othman, M. A. (2012). Short-term late-generation antibiotics versus longerterm penicillin for acute streptococcal pharyngitisinchildren. *TheCochraneLibrary*.
- 81. vandenAnker, J.N.(2013). Optimising the management of fever and pain in children. *International Journal of Clinical Practice*, 67(s178): 26-32.
- 82. Hay, A.D., Redmond, N.M., Costelloe, C., Montgomery, A. A., Fletcher, M., Hollinghurst, S., & Peters, T. J. (2009). Paracetamol and ibuprofen for the treatment of fever in children: the PITCHrandomisedcontrolledtrial. *Health Technol Assess*, *13*(27): 1-163.
- 83. Forrest, J.B., Heitlinger, E.L., & Revell, S.(1997). Ketorolacforpostoperativepain management in children. *Drug safety*, *16*(5): 309-329.
- 84. Arencibia, Z. B., & Choonara, I. (2012). Balancingthe risks and benefits of the use of

- over-the-counterpain medications in children. *Drug safety*, 35(12): 1119-1125.
- 85. Hayward, G., Thompson, M.J., Perera, R., DelMar, C. B., Glasziou, P. P., & Heneghan, C. J.(2012). Corticosteroids for the commoncold. *CochraneDatabaseSystRev*, 8(8).
- 86. Hayward, G., Thompson, M.J., Perera, R., Glasziou, P. P., Del Mar, C. B., & Heneghan, C. J.(2012). Corticosteroidsasstandaloneoradd-ontreatmentforsorethroat. *TheCochraneLibrary*.
- 87. Cingi, C., Songu, M., Ural, A., Erdogmus, N., Yildirim, M., Cakli, H., & Bal, C. (2011). Effect of chlorhexidinegluconateandbenzydaminehydrochloride mouth spray on clinical signs and quality of life of patients with streptococcaltonsillopharyngitis: multicentre, prospective, randomised, double-blinded, placebo-controlledstudy. *The Journal of Laryngology & Otology*, 125(6): 620-625.
- 88. Ciuman, R.R. (2012). Phytotherapeuticandnaturopathic adjuvant therapies inotorhinolaryngology. *European Archives of Oto-Rhino-Laryngology*, 269(2): 389-397.
- 89. Walton, J., Ebner, Y., Stewart, M.G., & April, M. M. (2012). System aticreview of randomized controlled trials comparing intracapsular tonsillec to my with to taltonsillecto my inapediatric 012). Management of a cute pharyngit is in children: summary of the Italian National Institute of Health guide lines. *Clinicaltherapeutics*, 34(6): 1442-1458.