Review Article

Actiology and the use of Antibiotics in the Case of Acute Pharyngitis: A Review

Kannan I¹*, Beulah Edwin¹, Vincent Prasanna², Hemlata Katiyar VM², Elango Dhanapal³

¹Department of Microbiology, Tagore Medical College and Hospital, Rathinamangalam, Chennai -600 127, India ²Department of ENT, Tagore Medical College and Hospital, Rathinamangalam, Chennai -600 127, India

Available Online: 01st May, 2015

ABSTRACT

Acute pharyngitis in one of the most common illness reported in high rate in an outpatient setup. The acute pharyngitis is caused due to infection by virus or bacteria. It is no clinical evidence that the bacterial illness and viral illness differ in their severity and duration. Hence the clinicians are put in difficulty in the selection of treatment modality especially the administration of antibiotics. However many research have proved that the adult pharyngitis is mainly due to virus. This review gives the picture of the aetiology and current treatment scenario of the acute pharyngitis.

Keywords: Acute pharyngitis, virus, bacteria, treatment, antibiotics

INTRODUCTION

Acute pharyngitis in one of the most common illness reported in high rate in an outpatient setup¹. The common symptoms include fever, sore throat and pain on swallowing. It can cause extreme distress of the body and can affect the routine activities of the patient. The illness is of great concern to the clinicians as it can easily spread to others if patient is present in close quarters. Even today the management of acute pharyngitis remains challenging to the clinicians because of its varied aetiology². Further the signs and symptom of the disease varies from patient to patient and thus makes the clinician difficult to ascertain proper aetiology of the illness. Even as the clinical diagnosis fails in many cases, the laboratory diagnosis of the disease also many times not reliable and may be misleading. The acute pharyngitis is the inflammation of soft tissues of the throat and can arise in many upper respiratory tract infections. Thus the symptoms can be presented in various degrees and thus cannot help in clinical diagnosis. Thus acute pharyngitis still remains as a challenge to the clinicians³.

AETIOLOGY

The acute pharyngitis is caused due to infection by virus or bacteria⁴. It is no clinical evidence that the bacterial illness and viral illness differ in their severity and duration⁵.

Viral pharyngitis

The viruses are considered to play a major role in the pathogenesis of pharyngitis⁶. It was found that viral pharyngitis account for around 70% of all pharyngitis, with bacterial causing only 20% to 40% of pharyngitis⁷. The Table 1 gives the possible viruses that are involved in the acute pharyngitis¹.

Those respiratory viruses, such as adenovirus, rhinovirus, respiratory syncytial virus, influenza virus and parainfluenza virus are the main cause of acute pharyngitis. The role of coxsakeivirus, herpesimplex virus and echoviruses in acute pharyngitis is not ruled out. The Epstein - Barr virus is also involved in acute pharyngitis, however is often accompanied with other clinical features such as generalized lymphadenopathy and splenomegaly that are characteristic of infectious mononucleosis. Certain viruses that cause systemic infections like rubella virus, measles and other viruses are also associated with acute pharyngitis. Eventhough viral acute pharyngitis is common in adults, many studies have revealed the role of viruses, especially adenovirus and Respiratory syncicial virus in acute childhood pharyngitis⁸⁻¹⁰.

Some viruses produces characteristic clinical symptom along with the pharyngitis. Adenoviruses can produce pharyngoconjunctival fever or an influenza-like syndrome known as the acute respiratory disease of military recruits¹¹. Coxsackieviruses are the most frequent causes of handfoot- and-mouth disease and herpangina¹². Many studies have revealed that the primary human herpesvirus 1 infection as a cause of pharyngitis^{13,14}. Human herpesvirus 2 can occasionally cause a similar illness as a consequence of oral–genital sexual contact¹⁵ *Bacterial pharyngitis*

Many studies have revealed that Group A streptococcus is by far the most common bacterial cause of acute pharyngitis, accounting for approximately 15 to 30 percent of cases in children and 5 to 10 percent of cases in adults^{16,17}. Apart from Group A streptococcus (GAS), some other bacteria are also have been demonstrated to be the aetiological agent of pharyngitis. Next to GAS, it

Virus	Symptoms/Disorders
Rhinovirus	Common cold
Coronavirus	Common cold
Adenovirus	Pharyngoconjunctival fever and acute respiratory disease.
Herpes simplex virus	Gingivostomatitis
types 1 and 2	
Parainfluenza virus	Cold and croup
Coxsakievirus A	Herpangina and hand – foot – and – mouth disease
Epstein – Barr virus	Infectious mononucleosis
Cytomegalovirus	Cytomegalovirus mononucleosis
HIV	Primary HIV infection
Influenza A and B viruses	Influenza

Table 1: Viruses that cause acute pharyngitis

has been shown the *Mycoplasma pneumoniae* and *Chlamydia pneumoniae* can also be involved in acute pharyngitis¹⁸. However, it has not been proved whether these bacteria are co-pathogens or the primary aetiological agents^{19,20}. In another study, apart from these bacteria, they have also found that *Legionella pneumophila* also can be the causative agent of pharyngitis²¹. Another important bacterium that has been implicated in the causation of pharyngitis is *Fusobacterium necrophorum*²². It occurs in patients aged 15 to 30 years. *F. necrophorum* can cause a severe complication, the Lemierre syndrome²³. This bacterium has now been emerged as an important bacterium in adolescent pharyngitis²⁴. It is considered to be as common as GAS in this age group²⁵.

In an interesting finding, the group C streptococcus has been found as the causative agent in acute pharyngitis has been frequently isolated from throat culture. Streptococcus *equi* subsp. *equisimilis* and *S. anginosus* are the two group C streptococci that have been isolate from pharyngitis²⁶. However, its role in the causation of pharyngitis is questionable^{27,28}. Certain studies have proved that it can cause pharyngitis²⁹⁻³².

Arcanobacterium haemolyticum is another bacterium which is rarely diagnosed to cause acute pharyngitis and tonsillitis in adolescents and young adults. The symptoms of infection caused by this organism closely mimic those of acute streptococcal pharyngitis, including a scarlatiniform rash in many patients^{33,34}. Normally the colonisation of the pharynx with *Neisseria gonorrhoeae* is asymptomatic, however it can occasionally cause pharyngitis³⁵.

TREATMENT

The multiple aetiology of acute pharyngitis makes it difficult for the physician to decide and initiate appropriate treatment for the patient. Treatment for acute pharyngitis nowadays more relies on patient satisfaction rather than the aetiology of the disease³⁶. However it is important for the physician to decide to initiate antibacterial therapy as in many cases the disease may be of viral aetiology. The physician should prescribe

Table 2a: Determination of patient's total sore throat score by assigning points to the following criteria (step 1)

1)	
Criteria	Points
Temperature above 38 ^o C	1
No cough	1
Tendor anterior cervical adenopathy	1
Tonsillar swelling or exudates	1
Age 3-14 years	1
Age 15-44 years	0
Age 45 years	-1

Table 2b: Total score calculated according to the above and choose the appropriate management suggested below according to the total sore throat score (Step 2)

Total score	Suggested management
0	No culture or antibiotic is required
1	
2	Culture all. Treat with antibiotics only if
3	culture result is positive
4	Culture all. Treat with penicillin on
	clinical grounds.

antibiotics in the suspected streptococcal pharyngitis to prevent the post complication sequels like peritonsillar or retropharyngeal abscess, cervical lymphadenitis, mastoiditis, sinusitis, and otitis media rheumatic fever and glomerulonephritis^{37,38}. However certain studies have shown that the post complications occur extremely rare even in the absence of antibiotic therapy³⁹. Furthermore, no evidence shows that antibiotic therapy for pharyngitis decreases the incidence of this complication⁴⁰. It has been universally accepted that the penicillin is the first choice treatment of streptococcal pharyngitis since GAS remains universally susceptible to penicillin^{41,42}. However ampicillin or amoxicillin are equally effective and thus can be given in place of penicillin⁴³. The studies have shown that treatment for 10 days with a single daily dose of amoxicillin is as effective as treatment with multiple daily doses of penicillin V^{44,45}. Amoxicillin became the antibiotic of choice in the place of penicillin in many countries as the penicillin is not commercially available. The patients who are allergic to beta-lactam antibiotics are administrated with macrolides^{46,47}. If the allergic patients, tested to have hypersensitivity to penicillin are not type I, cephalosporins should be considered as the alternative drug. Some studies have shown that cephalosporins has an efficacy higher than penicillin on GAS⁴⁸. Further cephalosporins have been found to be more effective than penicillin in case of recurrent streptococcal pharyngitis⁴⁹. The initiation of antibiotics has showed to be based on the clinical evidence of streptococcal pharyngitis. Patients with streptococcal pharyngitis commonly present with sore throat (generally of sudden onset), severe pain on swallowing, and fever. Headache, nausea, vomiting, and abdominal pain may also be present, especially in children⁵⁰. Thus the clinicians should clearly ascertain the possibility of streptococcal pharyngitis in a patient before initiating the antibiotic therapy. Some clinicians suggest the use of centor score to identify and treat the cases of pharyngitis^{51,52}. The Table 2 shows the calculation of centor score and the treatment approach according to it^{53,54}. The rapid antigen tests for identification of beta haemolytic streptococci have been used in certain hospital setups. When compared with the "golden standard" of throat culture, have reported sensitivities of 65% to 91% and specificities of 62% to 97%⁵⁵⁻⁵⁸. As these tests have approximately the sensitivity and greater specificity as that of throat culture, it can be used widely in all clinics to ascertain the cause of pharyngitis and thus can initiate the appropriate treatment⁵⁹.

CONCLUSION

To conclude, the indiscriminate use of antibiotics especially for adults in not advisable as from the review it is clear that the pharyngitis in adults are mainly due to virus. In a clinics a diagnostic and therapeutic rationale should be created to limit the use of antibiotic treatment to patients.

REFERENCES

- 1. Bisno, AL. Acute pharyngitis. N Engl J Med 2001; 344: 205–211.
- Susan M, Jutta P, Gregory J, Taj J, Deirdre C. Evaluation of potential factors contributing to microbiological treatment failure in *Streptococcus pyogenes* pharyngitis, Can. J. Infect. Dis. 2001; 12(1): 33-39.
- 3. Poses RM, Cebul RD, Collins M, Fager SS. The accuracy of experienced physicians' probability estimates for patients with sore throats: implications for decision making. JAMA, 1985; 254: 925-929.
- Kljakovic M. Sore Throat Presentation and management in general practice. N. Z. Med. J. 1993; 106: 381-383.
- Bisno, AL, Gerber MA, Gwaltney JM Jr, Kaplan EL, Schwartz RH. Diagnosis and management of group A streptococcal pharyngitis: a practice guideline. Clin Infect Dis 1997; 25, 574–583.
- Edwin B, Prasanna V, Kannan I, Katiyar VMH, Dhanapal E. Incidence of bacterial colonization in the oropharynx of patients with ear, nose and throat infections. Int J Med Sci Public Health. 2014; 3(8): 931-934.
- Stillerman M, Bernstein SH. Streptococcal pharyngitis. Evaluation of clinical syndromes in diagnosis. Am. J. Dis. Child, 1961; 101:476-489.
- 8. Putto A. Febrile exudative tonsillitis: viral or streptococcal? Pediatrics 1987; 80: 6–12.
- Putto A, Meurman O, Ruuskanen O. C-reactive protein in the differentiation of adenoviral, Epstein– Barr viral and streptococcal tonsillitis in children. Eur J Pediatr 1986; 145: 204–206.
- 10. Tsai HP, Kuo PH, Liu CC, Wang JR. Respiratory viral infections among pediatric inpatients and outpatients in Taiwan from 1997 to 1999. J Clin Microbiol 2001; 39, 111–118.
- 11. Hendrix RM, Lindner JL, Benton FR, et al. Large, persistent epidemic of adenovirus type 4-associated

acute respiratory disease in U.S. Army trainees. Emerg Infect Dis 1999; 5: 798-801.

- 12. Read RC. Orocervical and esophageal infection. In: Armstrong D, Cohen J, eds. Infectious diseases. Section 2. London: Harcourt, 1999; 33: 1-10.
- 13. McMillan JA, Weiner LB, Higgins AM, Lamparella VJ. Pharyngitis associated with herpes simplex virus in college students. Pediatr Infect Dis J 1993; 12: 280-4.
- 14. Glezen WP, Fernald GW, Lohr JA. Acute respiratory disease of university students with special reference to the etiologic role of *Herpesvirus hominis*. Am J Epidemiol 1975; 101: 111-21
- 15. Young EJ, Vainrub B, Musher DM, et al. Acute pharyngotonsillitis caused by herpesvirus type 2. JAMA 1978; 239: 1885-6
- 16. Poses RM, Cebul RD, Collins M, Fager SS. The accuracy of experienced physicians' probability estimates for patients with sore throats: implications for decision making. JAMA 1985; 254: 925-9.
- 17. Komaroff AL, Pass TM, Aronson MD, et al. The prediction of streptococcal pharyngitis in adults. J Gen Intern Med 1986; 1: 1-7.
- Principi N, Esposito S. Emerging role of Mycoplasma pneumoniae and Chlamydia pneumoniae in paediatric respiratory-tract infections. Lancet Infect Dis 2001; 1: 334–344.
- 19. Hammerschlag MR. The role of Chlamydia in upper respiratory tract infections. Curr Infect Dis Rep 2000; 2: 115–120.
- 20. Hammerschlag MR. Mycoplasma pneumoniae infections. Curr Opin Infect Dis 2001; 14: 181–186.
- 21. File TM Jr, Tan JS, Plouffe JF. The role of atypical pathogens: Mycoplasma pneumoniae, Chlamydia pneumoniae, and Legionella pneumophila in respiratory infection. Infect Dis Clin North Am 1998; 12: 569–592.
- 22. Amess JA, O'Neill W, Giollariabhaigh CN, Dytrych JK. A six-month audit of the isolation of Fusobacterium necrophorum from patients with sore throat in a district general hospital. Br J Biomed Sci. 2007; 64: 63-5.
- 23. Batty A, Wren MW. Prevalence of *Fusobacterium necrophorum* and other upper respiratory tract pathogens isolated from throat swabs. Br J Biomed Sci. 2005; 62: 66-70.
- 24. Jensen A, Hagelskjaer Kristensen L, Prag J. Detection of *Fusobacterium necrophorum* subsp. *funduliforme* in tonsillitis in young adults by real-time PCR. Clin Microbiol Infect. 2007; 13: 695-701.
- 25. Centor RM. Expand the pharyngitis paradigm for adolescents and young adults. Ann Intern Med 2009 Dec 1; 151: 812.
- 26. Fox, K., J. Turner, and A. Fox. 1993. Role of betahemolytic group C streptococci in pharyngitis: incidence and biochemical characteristics of *Streptococcus equisimilis* and *Streptococcus anginosus* in patients and healthy controls. J. Clin. Microbiol. 1993; 31:804–807.

- 27. Cimolai, N., R. W. Elford, L. Bryan, C. Anand, and P. Berger. 1988. Do the beta-hemolytic non-group A streptococci cause pharyngitis? Rev. Infect. Dis. 1988; 10: 587–601.
- 28. Hayden GF, Murphy TF, Hendley JO. Non-group A streptococci in the pharynx. Pathogens or innocent bystanders? Am. J. Dis. Child. 1989; 143: 794–797.
- 29. Meier F, Centor R, Graham L, Dalton H. Clinical and Microbiological Evidence for Endemic Pharyngitis Among Adults due to Group C Streptococci. Arch Intern Med. 1990; 150: 825-9
- 30. Turner J, Hayden F, Lobo M, Ramirez C, Murren D. Epidemiologic Evidence for Lancefield Group C Beta-hemolytic Streptococci as a Cause of Exudative Pharyngitis in College Students. J Clin Microbiol. 1997; 35: 1–4.
- 31.Jeffrey Tiemstra, Rosita LF. Miranda Role of nongroup A streptococci in acute pharyngitis Journal of the American Board of Family Medicine. 2009; 22(6): 663-669.
- 32. Al-Charrakh AH, Al-Khafaji JKT, Al-Rubaye RHS. Prevalence of -hemolytic groups C and F streptococci in patients with acute pharyngitis. North Am J Med Sci 2011; 3: 129-136.
- *33.* Karpathios T, Drakonaki S, Zervoudaki A, et al. *Arcanobacterium haemolyticum* in children with presumed streptococcal pharyngotonsillitis or scarlet fever. J Pediatr 1992; 121: 735-7.
- 34. Miller RA, Brancato F, Holmes KK. *Corynebacterium hemolyticum* as a cause of pharyngitis and scarlatiniform rash in young adults. Ann Intern Med 1986; 105: 867-72.
- 35. Wiesner PJ, Tronca E, Bonin P, Pederson AHB, Holmes KK. Clinical spectrum of pharyngeal gonococcal infection. N Engl J Med 1973; 288: 181-185.
- 36. Little P, Williamson I, Warner G, Gould C, Gantley M, Kinmonth AL. Open randomised trial of prescribing strategies in managing sore throat. BMJ. 1997; 314: 722-727.
- 37. Krober MS, Bass JW, Michels GN. Streptococcal pharyngitis: placebo-controlled double-blind evaluation of clinical response to penicillin therapy. JAMA 1985; 253: 1271-1274.
- Randolph MF, Gerber MA, DeMeo KK, Wright L. Effect of antibiotic therapy on the clinical course of streptococcal pharyngitis. J Pediatr 1985; 106: 870-875.
- 39. Chamovitz R, Catanzaro FJ, Stetson CA, Rammelkamp CH. Prevention of rheumatic fever by treatment of previous streptococcal infections. N Engl J Med. 1954; 251: 466-71.
- 40. Goslings WR, Valkenburg HA, Bots AW, Lorrier JC. Attack rates of streptococcal pharyngitis, rheumatic fever and glomerulonephritis in the general population. N Engl J Med. 1963; 268: 687-94.
- 41. Schwartz RH, Wientzen RL Jr, Pedreira F, Feroli EJ, Mella GW, Guandolo VL. Penicillin V for group A streptococcal pharyngotonsillitis: a randomized trial of seven vs ten days' therapy. JAMA 1981; 246: 1790-5.

- 42. Kaplan EL, Johnson DR, Del Rosario MC, Horn DL. Susceptibility of group A beta-hemolytic streptococci to thirteen antibiotics: examination of 301 strains isolated in the United States between 1994 and 1997. Pediatr Infect Dis J. 1999; 18: 1069-72
- 43. Choby BA: Diagnosis and treatment of streptococcal pharyngitis. Am Fam Physician 2009, 79:383-90.
- 44. Feder HMJ, Gerber MA, Randolph MF, Stelmach PS, Kaplan EL. Once-daily therapy for streptococcal pharyngitis with amoxicillin. Pediatrics 1999; 103: 47-51.
- 45. Shvartzman P, Tabenkin H, Rosentzwaig A, Dolginov F. Treatment of streptococcal pharyngitis with amoxycillin once a day. BMJ 1993; 306:1170–2.
- 46. Principi N, Esposito S. Comparative tolerability of erythromycin and newer macrolide antibacterials in paediatric patients. Drug Saf 1999; 20: 25–41.
- 47. Casey JR, Pichichero ME. Meta-analysis of cephalosporin versus penicillin treatment of group A streptococcal tonsillopharyngitis in children. Pediatrics 2004; 113:866-82.
- 48. Casey JR, Pichichero ME. The evidence base for cephalosporin superiority over penicillin in streptococcal pharyngitis. Diagn Microbiol Infect Dis 2007;57:39-45.
- 49. Casey JR, Pichichero ME. Symptomatic relapse of group A beta-hemolytic streptococcal tonsillopharyngitis in children. Clin Pediatr (Phila) 2007; 46(4):307-10.
- 50. Wannamaker LW. Perplexity and precision in the diagnosis of streptococcal pharyngitis. Am J Dis Child 1972; 124:352–8.
- 51. Snow V, Mottur-Pilson C, Cooper RJ, Hoffman JR. American Academy of family physicians; American College of Physicians-American Society of Internal Medicine; Centers for disease control. Principles of appropriate antibiotic use pharyngitis in adult. Ann Intern Med 2001; 134:506-8.
- 52. Centor RM, Allison JJ, Cohen SJ. Pharyngitis management: defining the controversy. J Gen Intern Med 2007; 22:127-30.
- 53. Centor RM, Witherspoon JM, Dalton HP, Brody CE, Link K. The diagnosis of strep throat in adults in the emergency room. Med Decis Making 1981; 1:239-46.
- 54. McIsaac W, Goel V, To T, Donald EL. The validity of a sore throat score in family practice. CMAJ 2000;163:811–15.
- 55. Dagnelie CF, Bartelink ML, van der Graaf Y, Goessens W, de Melker RA. Towards a better diagnosis of throat infections (with group A betahaemolytic streptococcus) in general practice. Br J Gen Pract. 1998;48:959-6
- 56. Roddey OF Jr, Clegg HW, Martin ES, Swetenburg RL, Koonce EW. Comparison of an optical immunoassay technique with two culture methods for the detection of group A streptococci in a pediatric office. J Pediatr. 1995; 126: 931-3.
- 57. Schlager TA, Hayden GA, Woods WA, Dudley SM, Hendley JO. Optical immunoassay for rapid detection of group A beta-hemolytic streptococci. Should

culture be replaced? Arch Pediatr Adolesc Med. 1996; 150:245-8.

58. Hart AP, Buck LL, Morgan S, Saverio S, McLaughlin JC. A comparison of the BioStar Strep A OIA rapid antigen assay, group A Selective Strep Agar (ssA), and Todd-Hewitt broth cultures for the detection of

group A Streptococcus in an outpatient family practice setting. Diagn Microbiol Infect Dis. 1997;29:139-45

59. Gerber MA, Tanz RR, Kabat W, Dennis E, Bell GL, Kaplan EL, et al. Optical immunoassay test for group A beta-hemolytic streptococcal pharyngitis. An officebased, multicenter investigation. JAMA. 1997;277:899-903