ORIGINAL ARTICLE

Importance of High C-Reactive Protein (CRP) And Total Leukocyte Count (TLC) In Odontogenic Space Infections

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ABSTRACT

Background: An acute inflammatory homo-pentameric plasma protein, C-reactive Protein, discovered in 1930, is found in small amounts in healthy persons and plays an important role in natural immune system, activation of compliment system and phagocytosis and antigen clearance.

Aim: To determine high C - reactive protein (CRP) and Total Leukocyte Count (TLC) in patients suffering from odontogenic space infections.

Methods: Present study was carried out in the department of Oral Surgery at Islam Dental College, Sialkot from Sept. 2020 to Oct 2021. A total of 151 male and female patients of 22-65 years age range with odontogenic space infection were included. Patients suffering from chronic disease like diabetes, chronic renal failure, and pregnant women were excluded. Sample of patient's blood was sent to the hospital laboratory for measurement of serum C - reactive protein (CRP) and Total Leukocyte Count (TLC) levels.

Results: Patient's age range in our study was 22 to 65 years with mean age of 54.5 ± 14.68 years. Out of 151 patients, 93 (61.58%) were male and females were 58 (38.42%) showing male to female ratio of 1.6:1. Mean CRP levels were 3.52 ± 1.23 mg/L and Mean TLC levels were 13670 ± 1890 cells/mm3. In our study, frequency of raised CRP levels was found in 100% and TLC in 71.82% of patients with odontogenic space infections.

Conclusion: This study concluded that frequency of raised C-Reactive Protein (CRP) levels are found in 100% patients, indicating CRP to be an effective biomarker in infected patients.

Keywords: Odontogenic Infections, C-Reactive Protein (CRP), Total Leukocyte Count (TLC).

INTRODUCTION

Most of orofacial infections in human beings are odontogenic in origin.¹ Patients presenting odontogenic space infections of the head and neck region are at more risk for life threatening situations due to minimal resistance, anatomical location, vital organ's presence in the area and fast spread mode of these infections.² Several Serious complications of Respiratory and cardiovascular system may become inevitable; therefore, careful scrutiny and monitoring of these patients is necessary. Such serious complications usually occur at a rate of 10-20%^{3,4}. A serum inflammatory biomarker like high sensitivity C-reactive protein (hs-CRP), has been advocated to highlight the risk of adverse cardiovascular (CV) events effectively⁵⁻¹¹.

cardiovascular (CV) events effectively⁵⁻¹¹. White Blood Count (WBC) count and Erythrocyte Sedimentation Rate (ESR) values are important at testing time, however their predictability is limited.¹² Due to importance of serum-derived surrogate predictor behavior and outcome, several inflammatory markers are discovered functioning as prospective indicator of disease process. ¹³ in 1930, Tillet and Francis discovered C-reactive protein (CRP) during the investigatin of sera of Pneumococcus infectious patients and named it due to its (C)-polysaccharide reaction with the capsular of *Pneumococcus*^{14,17, 22} C-reactive Protein is found in small amounts in healthy persons and plays an important role in natural immune system, activation of compliment system and phagocytosis and thus antigen clearance.^{15,16} Marked increase up to 1000 fold of serum CRP levels, is noted within a few hours of clinical symptoms in severe inflammatory reactions.¹⁶ In a study conducted by Bali R and colleagues in 2017, raised CRP levels were found in 100% and TLC in 64% of patients with Odontogenic space infections^{17,18}.

Received on 11-09-2021 Accepted on 22-03-2022 Total Leukocyte Count (TLC) test is commonly used for diagnosing Odontogenic space infections, adding the estimation of CRP as a new diagnostic tool.

This study was conducted to determine raised CRP and TLC in odontogenic space infection patients. Literature search revealed no local study on this topic.

MATERIALS AND METHODS

This descriptive, cross-sectional study was done after IRB permission in Oral Surgery Department at Islam Dental College Sialkot. Total of 151 patients of 22-65 years age range, were diagnosed with odontogenic space infections by taking patient's history, thorough clinical examination and radiographic Examination indicating periapical changes. We excluded patients suffering from chronic disease like diabetes mellitus, chronic renal failure and pregnant women from the study. The study was approved by the ethical review committee of Islam Dental College, Sialkot

An informed patient's consent and college ethical committee permission was taken. Collected Samples of patient's blood for serum C - reactive protein (CRP) and Total Leukocyte Count (TLC) level measurements were sent to hospital laboratory. The results were analyzed by SPSS version 20.0 and Mean and standard deviation for age, duration of symptoms and raised serum C-Reactive Protein (CRP) and Total Leukocyte Count (TLC) levels, their raised frequency and percentages were calculated for gender.

RESULTS

Age range of patients in our study was 22-65 years with mean of 54.5 \pm 14.68 years. Out of 151 patients, 93(61.58%) were male and 58 (38.42%) were females and male to female ratio was 1.6:1. Mean duration of symptoms in our study was 29.24 \pm 7.61 hours. Mean BMI was 30.02 \pm 2.05kg/m². Mean CRP levels were 3.52 \pm 1.23 mg/L. Mean TLC levels were 13670 \pm 1890 cells/mm³. In

Table 1: Raised C-Reactive Protein (CRP) and Total Leukocyte Count (TLC) in patients with odontogenic space infections

Laboratory Test	Frequency (%)	
	Yes	No
Raised CRP in Patients	151(100%)	0
Raised TLC in patients	108 (71.82%)	43 (28.18%)



DISCUSSION

According to many authors, Inflammatory serum markers are very useful and quantitatively can be helpful for finding therapeutic efficacy of various treatment modalities for investigating the severity of infection, dealing with pre and post-operative infections and appropriate use of prophylactic antibiotic. ^{20,21} Thus many authors advised use of CRP in infection assessment due to their multiple advantages^{20,18}.

This study was conducted to find the frequency of raised C reactive protein and Total Leukocyte Count (TLC) in odontogenic space infections. Age of patients in this study ranged between 22 to 65 years with mean of 54.5 ± 14.68 years. Out of 151 patients, male were 93(61.58%) and females were 58(38.42%) with 1.6.1 male to female ratio. In our study, frequency of raised CRP levels was found in 100% and TLC in 71.82% patients with Odontogenic space infections. In a study conducted by Bali R and colleagues in 2017, raised CRP levels were found in 100% and TLC in 64.0%patients with Odontogenic space infections¹⁷.

Tillet and Francis identified CRP in 1930 as an acute-phase protein²². It has been studied as a screening device for inflammation, a non-specific marker for disease activity and as a diagnostic adjunct. Physiologically, it increases cell-mediated immunity by phagocytosis promotion, chemotaxis acceleration and platelets activation. CRP is a reliable early indicator of inflammation or injury^{22,23}. According to Mustard et al. CRP post-op levels could indicate septic complications prior to their clinical appearance²⁴.

In similar studies by Pinilla et al., and Malve I found statistically significant correlation between pre-albumin and CRP at 2nd day (r =0.45, p < 0.01) and 5th day (r =0.53, p< 0.01) and mean serum levels of pre-albumin decreased (p < 0.001) and significantly increased levels of CRP (p < 0.01) respectively in patients compared to healthy controls^{25,17} according to studies by Pinilla et al²⁷, use of Pre-albumin and CRP inflammatory markers is an easy and inexpensive method of assessment of illness

severity in critical patients²⁷. CRP can be a useful infection indicator along higher body temperature above 38.2°C also stated by Povoa et al in a study where healthy individual's CRP raised from 0.08 mg/dl to more than 8.7mg/dl when infected by S aureus⁴⁰.

Importance of WBCs is more in the assessment of patient's response to treatment but has minor role in the diagnosis and judgement of severity of head and neck infections which is well supported by studies of Boucher et al²⁹ and Heimdahl and Nord ³⁰. When Compared in odontogenic infections, CRP is better marker in an infection process due to faster increase in level of CRP than WBC^{31,32}. In a study of comparative assessment, Ebersole in 1997³³, also found significant raised CRP and hepatoglobin in periodontitis patients and normal person's characteristics. With this finding Manelli in 1998 and others came to conclusion that localized infection and inflammation, resulting in stimulating systemic host changes are manifested by raised CRP^{34 35}.

Similarly CRP, IL-6 and α -1 antitrypsin concentration levels were found raised in the serum in a study by Gunnel et al. and Haupt et al^{36,37,38}. Flores et al. in 2001 also concluded higher CRP levels suggesting underlying infectious complication while studying several patients with severe infection due to trauma.³⁹ Therefore, raised CRP levels show an infectious condition of the body, which in case of our study are the odontogenic spaces.

The limitation of this study was that a relatively smaller sample size was used and there was no long term follow up. Further studies with longer follow ups and a larger sample size are recommended so that the results can be generalized to the local population.

CONCLUSION

This study concluded that frequency of raised CRP concentration was found in 100% patients, and CRP was found to be an effective biomarker in patients with odontogenic space infections as compared to TLC. So, we recommend that CRP should be used in assessment and management of odontogenic space infections routinely to reduce the morbidity of such patients.

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REFERENCES

- Naik S, Balreddy D, Sridhar D, Prasad D. Efficacy of serum C reactive protein levels as monitoring tools for patients with infections of odontogenic origin - A clinical and biochemical study. Int J Healthcare Biomed Res. 2015;04:32-9.
- Bahl R, Sandhu S, Singh K, Sahai N, Gupta M. Odontogenic infections: microbiology and management. Contemp Clin Dent. 2014;5:307-11.
- Gujrathi AB, Ambulgekar V, Kathait P. Deep neck space infection a retrospective study of 270 cases at tertiary care center. World J Otorhinolaryngol Head Neck Surg. 2016;2:208-13.
- 4- Arslan Masood, Shariq Sohail Jafar, Zubair Akram. Serum high sensitivity C-reactive protein levels and the severity of coronary atherosclerosis assessed by angiographic gensini score. JPMA. April 2011, Volume 61, Issue 4.
- 5- Koenig W, Lowel H, Baumert J, Meisinger C. C-reactive protein modulates risk prediction based on the Framingham Score: implications for future risk assessment: results from a large cohort study in southern Germany. Circulation 2004; 109: 1349-53.
- 6- Ballantyne CM, Hoogeveen RC, Bang H, Coresh J, Folsom AR, Heiss G, et al. Lipoprotein-associated phospholipase A2, highsensitivity C-reactive protein, and risk for incident coronary heart disease in middle-aged men and women in the Atherosclerosis Risk in Communities (ARIC) study. Circulation 2004; 109: 837-42.
- 7. Danesh J, Wheeler JG, Hirschfield GM, Eda S, Eiriksdottir G, Rumley A, et al. C-reactive protein and other circulating markers of inflammation in the prediction of coronary heart disease. N Engl J Med 2004; 350: 1387-97.

- 8- Pai JK, Pischon T, Ma J, Manson JE, Hankinson SE, Joshipura K, et al. Inflammatory markers and the risk of coronary heart disease in men and women. N Engl J Med 2004; 351: 2599-610.
- 9- Cushman M, Arnold AM, Psaty BM, Manolio TA, Kuller LH, Burke GL, et al. C-reactive protein and the 10-year incidence of coronary heart disease in older men and women: the cardiovascular health study. Circulation 2005; 112: 25-31.
- 10-Ridker PM, Rifai N, Cook NR, Bradwin G, Buring JE. Non-HDL cholesterol, apolipoproteins A-I and B100, standard lipid measures, lipid ratios, and CRP as risk factors for cardiovascular disease in women. JAMA 2005; 294: 326-33.
- Laaksonen DE, Niskanen L, Nysskonen K, Punnonen K, Tuomainen TP, Salonen JT. C-reactive protein in the prediction of cardiovascular and overall mortality in middle-aged men: a population-based cohort study. Eur Heart J 2005; 26: 1783-9.
- Rastenienė R, Pūrienė A, Aleksejūnienė J, Pečiulienė V, Zaleckas L. Odontogenic maxillofacial infections: a ten-year retrospective analysis. Surg Infect (Larchmt). 2015;16:305-12.
- Goumenakis D, Giannakopoulos NN, Parara E, Mourouzis C, Rallis G. Effect of causative tooth extraction on clinical and biological parameters of odontogenic infection: a prospective clinical trial. J Oral Maxillofac Surg. 2015;73:1254-8.
- 14-Pal S, Pal GY, Yadu MFA, Yadu S. Efficacy of serum prealbumin as monitoring tools for patients with fascial space infections of odontogenic origin- a clinicobiochemical study. Int J SciEngin Res. 2015; 6:2144-8.
- Sharma A, Giraddi G, Krishnan G, Shahi AK. Efficacy of serum prealbumin and crplevels as monitoring tools for patients with fascial space infections of odontogenic origin: a clinicobiochemical study. J Maxillofac Oral Surg. 2014;13:1-9.
- Bagul R, Chandan S, Sane VD, Patil S, Yadav D. Comparative evaluation of C - reactive protein and WBC count in fascial space infections of odontogenic origin. J Maxillofac Oral Surg. 2017; 16:238-42.
- Bali R, Sharma P, Ghanghas P, Gupta N, Tiwari JD, Singh A, et al. To compare the efficacy of c-reactive protein and total leucocyte count as markers for monitoring the course of odontogenic space infections. J Maxillofac Oral Surg. 2017;16:322-7.
- 18- Nicola R. Sproston and Jason J. Ashworth. Role of C-Reactive Protein at Sites of Inflammation and Infection. Front Immunol. 2018; 9: 754
- Imran M, Shetty A, Vivek GK, Shafath A, Vaibhav N. C-Reactive Protein and its applications in Oral and Maxillofacial Surgery- An Overview. Saudi J Oral Dent Res. 2017;2:93-7.
- Sann L. Evolution of serum prealbumin, C- reactive protein and orosomucoid in neonates with bacterial infection. J Pediatric 1984:105: 977-81.
- 21. Ylyjoki S, Suuronen R, Somer HJ, Meurman JH, Lindquist C. Differences between patients with or without the need for intensive

care due to severe odontogenic infections. J Oral maxillofacial Surg 2001; 59:867 -72.

- Tillet WS, Francis T Jr. Serological Reactions in pneumonia with a non-protein somatic fraction of pneumococcus. J Exp Med. 1930; 52:561–71.
- Pepys MB, Baltz ML. Acute phase proteins with special reference to C - reactive protein and related proteins and serum amyloid A protein. AdvImmunol. 1983; 34:141–212.
- Zimmerman MA, Selzman CH, Cothren C, Sorensen AC, Raeburn CD, Harken AH. Diagnostic implications of C - reactive protein. Arch Surg. 2003; 138:220–4.
- Ridker PM. Prospective study of C- reactive protein and the risk of future cardiovascular events among apparently healthy women. Circulation. 1998; 98:731–3.
- Manelli JC, Badetti C, Golstein MM, Bernard D. A reference standard for plasma proteins is required for nutritional assessment of adult burn patients. Burns. 1998; 24:337–45.
- Pinilla JC. The C-reactive protein to prealbumin ratio correlates with the severity of multiple organ dysfunction. Surgery. 1998; 124:799– 806.
- Nagarale G, Ravindra S, Thakur S, Setty S. Efficacy of a chairside diagnostic test kit for estimation of C-reactive protein levels in periodontal disease. J Indian SocPeriodontol 2010;14:213–6.
- 29. Boucher NE, Hanharan JJ, Kihara FY. Occurrence of C-reactive protein in oral diseases. J Dent Res. 1967;46:624–7.
- Heimdahl A, Nord CE. Orofacial infections of odontogenic origin. Scand J Infect Dis. 1983;39(Suppl):86–91.
- Kallio LU, Kallio MJ, Peltola H, Eskola J. Serum C-reactive protein, erythrocyte sedimentation rate, and white blood cell count in acute hematogenous osteomyelitis of children. Pediatrics. 1994; 93:59–62.
- Clyne B, Olshaker JS. The C-reactive protein. J Emerg Med. 1999; 17:1019–25.
- Ebersole JL. Systemic acute phase protein reactants, C-reactive protein and hepatoglobin in adult periodontitis. Clin Explmmunol. 1997; 107:347–52.
- Lagrand WL. C-reactive protein as a cardiovascular risk factor. J Am Heart Assoc. 1999; 100:96–102.
- Yeun JY. C-reactive protein predicts all-cause and cardiovascular mortality in hemodialysis patients. Am J Kidney Dis. 2000; 35:469–76.
- Haupt W. Association between preoperative acute phase response and postoperative complications. Eur J Surg. 1997; 163:39–44.
- Flores JM. C reactive protein as a marker of infection among patients with severe closed trauma. Enferm Infect Microbiol Clin. 2001; 19:61– 5.
- Sullivan DH, Bopp MM, Roberson PK. Protein-energy under nutrition and life threatening complications among the hospitalized elderly. J Gen Intern Med. 2002; 17:923–32.
- Ortega O. Significance of C- reactive protein levels in pre-dialysis patients. Nephrol Dial Trans. 2002; 17:1105–9.
- Povoa P, Coelho L, Almeida E, Fernandes A, Mealha R, Moreira P, et al. C-reactive protein as a marker of infection in critically ill patients. *Clin Microbiol Infect* (2005) 11(2):101–8