

Evaluation of Interleukin IL-1 β , TNF- α , and VDB Protein Levels in Serum of Patients with Chronic Periodontitis

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Abstract

Aim: The aim of the study was to evaluate the levels of Interleukin IL-1 β , TNF α , and VDB Protein in the serum of patients with chronic periodontitis. **Materials and methods:** Serum samples of 60 persons, aged 25-55 years with chronic periodontitis were collected as patients group; samples of serum were obtained from 30 healthy subjects aged 25-55 years as the control group. Interleukin IL-1 β , TNF- α , and VDB protein levels were measured by enzyme-linked immune sorbent assay (ELISA) kit. **Results:** The levels of Interleukin IL-1 β , TNF- α , and VDB protein in serum showed a significant increase in the patient group than the healthy group. On the other hand, VDB protein was not significantly correlated with IL-1 β and TNF- α in patients.

Key words: Interleukin IL-1 β , TNF- α , VDB protein, chronic periodontitis

Introduction

Periodontal diseases can be defined as a group of inflammatory diseases that are influenced by host response factors. In the last years, there was a consensus toward the converse relationship between periodontal disease and systemic disease (Elkadi et al., 2018). periodontitis is divided into two types, aggressive periodontitis, and chronic periodontitis. Clinically, Chronic periodontitis is distinct from other less common periodontal diseases such as aggressive and necrotizing ulcerative periodontitis. The onset age of the chronic form is higher than the age for the aggressive forms (Martu et al., 2014). As a response to the periodontal pathogens and their endo-toxins, The immune cells in the periodontium release proinflammatory mediators (Liu et al., 2001).

TNF- α is a pro-inflammatory cytokine that plays an important role in the inflammatory reaction. There are two types of TNF, TNF- α and TNF- β . TNF- α is produced by the activated macrophages, neutrophils, keratinocytes, monocytes, and mast cells in response to lipopolysaccharides (LPSs) (Newman et al.,

2006). TNF- α has Many actions such as pro-inflammatory, facilitation of Leukocyte recruitment and vascular permeability, mediated Macrophage-induced angiogenesis, and vascular proliferation in the periodontal granulation tissue formation. In response to bacterial LPS, TNF- α triggers osteoclast activation, proliferation, and differentiation resulting in bone resorption (Varghese et al., 2015).

Interleukin (IL)-1 is a potent pro-inflammatory molecule produced by macrophages and marrow stromal cells and found in two active forms, IL-1 α and IL-1 β encoded by separate genes. Both are the main constituents of what was once called osteoclast-activating factor (Rosenvall, 2013). In addition to pro-inflammatory, the action of Interleukin (IL)-1 stimulates bone resorption and implicated in pathological conditions with bone loss (Stashenko et al., 1991). IL-1 β has been found to be significantly increased in the periodontal tissues compared with healthy sites (Afnan Abdulkareem Hussain BDS and Basima Ghafory, 2014).

Vitamin D binding protein (DBP) is a multifunctional protein, that acts as inflammatory regulation and is found in plasma. The major function of Vitamin D binding protein (DBP) is the ability to bind to vitamin D (calcitriol) and transfer it to target cells. Vitamin D was shown to be involved in both the innate and adaptive immune systems, with vitamin D insufficiency being linked to many inflammatory disorders, including periodontal diseases. It has been suggested that vitamin D may act similarly to cytokines, and regulate the inflammatory process by several mechanisms including stimulating phagocytosis and antibody presenting actions to enhance the initial immune response. As the inflammatory process progresses, vitamin D plays a role in the inhibition of T-cell proliferation and thus inflammatory resolution (Aboodi et al., 2016; Stein et al., 2014).

Materials and Methods

Sample collection

serum of 60 patients with chronic periodontitis from teaching hospital of College of Dentistry, University of Baghdad was collected at 10.00-12.00 AM. Another 30 healthy samples were collected. 5-ml of venous blood was collected from each patient and control. The blood samples were centrifuged for 10 minutes at 1500 rpm, and the separated serum sample was kept at -40 °C till used.

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Interleukin IL-1 β , TNF- α , and VDB protein assay

Interleukin 1L- β , TNF- α , and VDB protein concentrations were measured by enzyme-linked immune sorbent assay (ELISA) kit (My BioSource, USA).

Statistical analysis

Statistical analysis was done using SPSS version 10.0 and values were expressed as mean \pm SD. The comparison of mean \pm SD was performed using Student t-test and correlation analysis was performed using Pearson's correlation test. Statistical significance was defined as $P \leq 0.05$.

Results and Discussion

Interleukin IL-1 β , TNF- α , and VDB protein were expressed as mean \pm SD in the sera of chronic periodontitis patients compared to the normal group (Table 1).

Table 1: Interleukin 1L-1 β , TNF- α , and VDB protein levels (pg/ml) (mean \pm SD) in all patients compared to the normal group.

	Groups	No.	(mean \pm SD)	Probability
TNF- α	Control	30	237.976 \pm 55.519	0.001
	patients	60	318.479 \pm 78.650	
IL-1 β	controls	28	124.880 \pm 25.204	0.001
	patients	60	144.434 \pm 23.443	
VDB protein	controls	28	127.198 \pm 48.397	0.001
	patients	60	144.537 \pm 40.423	

The data in Table 1 shows that TNF- α levels were elevated significantly ($p < 0.05$) in the patient group compared with the control group. These findings were in agreement with the results of many studies such as Loana *et al.* (2014), Ravindra *et al.* (2012), and Thilagar *et al.* (2018) that found the TNF- α mean levels were higher in chronic periodontitis group compared to control group. Omneya *et al.* (2018) and sazan *et al.* (2016) found a significant increase in TNF- α level in patients with chronic periodontitis compared with the control group and observed a statistically significant reduction in TNF- α levels after 2 months from periodontal therapy. In contrast to our results, Yamazaki *et al.* (2005) found that TNF- α and IL-6 levels didn't change following periodontal treatment, and there was no difference in the serum levels of these inflammatory cytokines between patients and control subjects.

TNF- α is part of the major pro-inflammatory cytokines which are usually produced on inflammatory sites by the mononuclear cells infiltration. In consequence, this cytokine is part of the periodontal tissue destruction. Moreover, meningococcal induces the synthesis of IL-1 and PGE-2, activates the osteoclasts, inducing bone resorption (Oates *et al.*, 2002). The high levels of TNF- α are related to different pathologic conditions, like sepsis shock, cachexia (HIV, tuberculosis), autoimmune diseases, hepatitis, leukemia, myocardial infarction, transplant rejection,

multiple sclerosis, rheumatoid arthritis, and meningococcal sepsis (Gokul, 2012).

Interleukin IL-1 β is the main cytokine identified in chronic periodontal disease, with further influence on the activity of immune cells (Grover *et al.*, 2016). The interleukin (IL)-1 family of cytokines has a central role in triggering and perpetuating the immune and inflammatory responses (Grover *et al.*, 2016). Pro-inflammatory cytokines, such as interleukin 1 β , play an important role in the initiation and regulation of immune responses in periodontium (Grover *et al.*, 2016).

In this study, we found a significant elevation in serum IL-1 β level among chronic periodontitis patients in comparison to that of healthy control. These findings are consistent with some studies such as Afnan *et al.* (2014) that recorded a significant increase in interleukin 1 β level in patients with chronic periodontitis compared with the control group. Gani *et al.* (2009), Tälvan *et al.* (2017), Acharya *et al.* (2015), and Finoti *et al.* (2017) agreed with the results of this study. Gorska *et al.* (2003) in Poland found that the levels of IL-1 β were significantly higher in serum and gingival tissue biopsies in chronic periodontitis patients as compared to healthy control. Al-Ghurabei *et al.* (2012) found that the detection of higher levels of serum IL-1 β in the serum of patients with chronic periodontitis was consistent with the cytotoxins' role in inflammation and suggests that the serum IL-1 β may be a good marker of periodontal inflammation. In contrast to the present study, other studies revealed that there was a nonsignificant difference in serum IL-1 β concentration between patients with chronic periodontitis group and healthy group (Al-Ghurabei, 2012). Aboodi (2015) found that the levels of VDB protein were increased in periodontal patients compared to the healthy group. Dietrich *et al.* (2005) observed the increased VDB protein levels during chronic periodontitis because of an increased abundance of vitamin D during this phase; high vitamin D levels were suggested to reduce inflammation. Our observation of increased DBP levels in chronic periodontitis agreed with the results of this study. In correlation study, we found no significant relationship between VDB protein and IL-1 β and TNF- α ($P > 0.005$) as shown in Figures 1 and 2.

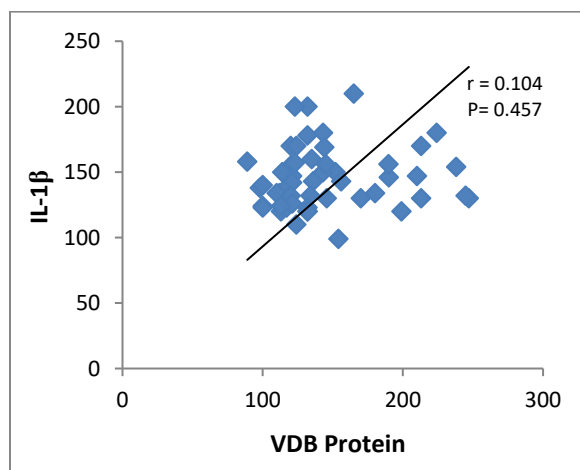


Figure 1: correlation between VDB Protein and IL-1 β .

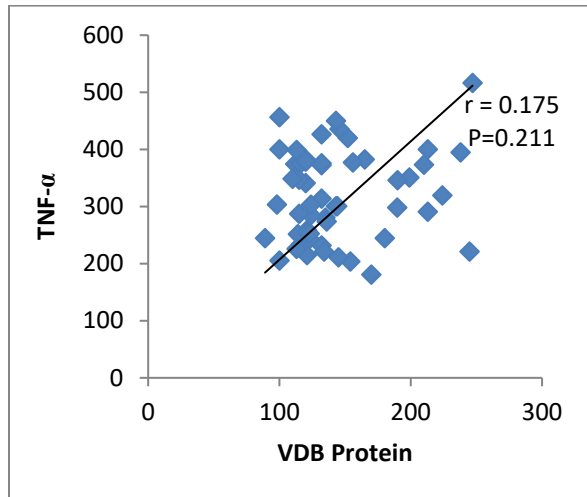


Figure 2: correlation between VDB protein and TNF- α .

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