RESEARCH ARTICLE

Procalcitonin as New Proinflammatory Marker in Burn Patients

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ABSTRACT

Procalcitonin (PCT) is a 116 amino acid peptide with a sequence identical to that of prohormone of calcitonin but devoid of hormonal activity. The aim of this study was to determine PCT and its use as a new prognostic pro-inflammatory marker in burn patients and determine the role of PCT level as a mortality indicator in survival and non-survival burn patients. The burn patients in this study were divided into four groups according to the percentage of total body surface area burned: carry out on (50) burn patients admitted to the burn unit in West Erbil Emergency Hospital in Erbil governorate and (20) apparently healthy non-burn individuals who regarded as healthy control group (HC) in this study from February 2012 to April 2012. Out of 50 burnt patients, 20 patients were second sampled to follow-up their immune profile. Regarding PCT there were significant differences ($P < 0.05$) observed between the mean serum concentration of PCT belong to 18 burn patients of 1st group (G1) (1.24 ± 0.82), 23 patients of 2nd group (G2) (1.52 ± 0.76), six patients of 3rd group (G3) (2.16 ± 0.11), and three burn patients of 4th group (G4) (6.5 ± 2.87) with mean serum concentration of 18 HC (0.02 ± 0.006). Significant differences obtained when compared to G3 (2.16 ± 0.11) versus G4 (6.5 ± 2.87) using the F-test and t-test. Results show that PCT is a highly productive laboratory parameter involving an easy and rapid bedside test for diagnosis and prognosis mortality in burn patients.

Keywords: Procalcitonin; Total body surface area; Burn patient

INTRODUCTION

The burn is an injury consisting of the destruction of the skin and the underlying tissues (Siamanga, 2002). It is a significant issue in the world and shows one of the most destructive trauma forms. More than 1,000,000 injuries of burn take place in the United States each year (Enkhbaatar, 2004). Burn can be caused by different agents, for example, heat (scald or flame), chemicals, electricity, freezing, friction, and radiation (Keck et al., 2009).

Various agents affect the mortality and morbidity of injury of burn. First, the burn size, based on the percent of the total body surface area (TBSA) is immediately proportionate to survival. Furthermore, the greater the risk of the systemic infection and subsequent burn wounds but the bigger the burn, which is directly related to rise morbidity and mortality (Sheridan, 2000). Second, the injury of inhalation in combination with burns worsens the clinical outcome of the burn patient to the great extent, despite advances in intensive respiratory support (Stern, 2009). Third, the patient age significantly affects outcomes after burns, as the mortality is higher in old and very young patients in comparison to other groups of age (Pham et al., 2009). It was exhibited time after time in the clinical investigations that mortality was highest in the old patient

who had smoke inhalation injury, or/and more severe burns and had finding implicit diseases (McGill et al., 2000).

The injury nature of burn wound is basically resulting of the complex processes causing local, besides systemic complications, affecting multiple organ systems distal to the skin (Evers et al., 2010). The pathogenesis of the progression of the burn wound starts from the original injury site. It includes loss of main tissue owing the denaturation of proteins, which further leads to releasing platelet activating factors and toxic pro-inflammatory assisters into the circulation (Piccolo et al., 1999). Involvement of the immune cell, such as macrophages and neutrophils, resulted in the producing the cytotoxic oxygen free-radical damaging the dermal structure indirectly, or directly (Singh et al., 2007).

Procalcitonin (PCT) is a 116 amino acid peptide with a sequence identical to that of prohormone of calcitonin but devoid of hormonal activity. Since the determination of PCT was first achieved by Assicot et al., 1993 for differentiating between nonbacterial and bacterial causatives of sepsis. One main merit of PCT compared with other parameters is its high and early particular increase in response to sepsis and severe systemic bacterial infections (Müller et al., 2000). Therefore, in the septic conditions

increased levels of PCT can be noticed 3–6 h after the challenge of infection. Therefore, the main aim of this study is to test PCT as a new prognostic pro-inflammatory marker in burn patients to indicate early and highly specific increases in response to bacterial infections and sepsis.

METHODS

Fifty burned patients who were admitted to West Erbil Emergency Hospital (WEEH) in Erbil governorate were included in this study from February 2012 to April 2012. Out of 50 burnt patients, 20 patients were second sampled to follow-up their immune profile. The PCT was estimated using (Human PCT ELISA kit) provided by RayBio/USA which is an *in vitro* enzyme-linked immunosorbent assay for the quantitative measurement of human PCT in serum with a normal range <0.5 ng/ml according to the manufacturer's specifications. The calculated TBSA performs by Lund and Browder chart. Blood samples were collected from burned patients, and transferred into 10 ml sterile tube and centrifuged at 3000 rpm for 3 min and serum was separate from whole blood and stored in deep freeze (-80°C) in aliquot into several Eppendorf tubes until assayed.

Ethical Consideration

The present work has been approved from the College of Medicine, Erbil at the research ethics committee of Hawler Medical University. All participants were obtained with written informed consent after procedures and purposes of the study were illustrated.

Experimental design

The burn patients in this study were divided into four groups: Group 1 (1–25%), Group 2 (26–50%), Group 3 (51–75%), and Group 4 (76–100%) according to the percentage of TBSA% burned carry out on (50) burn patients admitted to burn unit in WEEH in Erbil governorate and (20) apparently healthy non burn individuals who regarded as a healthy control group (HC). Out of 50 burnt patients, 20 patients were secondly sampled to follow-up their immune profile.

Statistical Analysis

Statistical analysis has been completed by Statistical Package for Social Sciences ver. 19. The relative proportions have been determined with a confidence interval of 95%. The possible association was identified by Fisher's exact and the Chi-square statistical tests at a probability of $<5\%$.

RESULTS

Distribution of burn patients according to gender showed that females represent the majority of burn trauma

32 (64%) in comparison to male were 18 patients (36%) with statistically no significant differences ($P > 0.05$), [Table 1].

On the basis of TBSA %, burned patients were categorized into four groups (G1, G2, G3, and G4). The highest number of burn injury located in the second group (G2) 23 (46%), with TBSA (26–50%), in which the female was higher than male as shown as in Table 2.

Out of 50 burn patients, 32 females and 18 males, the total number of mortalities was 18 (14 females and 4 males) as shown as in Table 3.

Concerning the survivors and non-survivors burn patients, the number of survivors burn patients in female was 18 (56.25%) and in male was 14 (43.75%) with mean TBSA% burned (25.23%) and mean age 20.79. Out of 18 non-survivors burn patient, a large number of burn victim fell in female in which 14 (77.78%), while in male was only 4 (22.22%) with mean TBSA % burned (52.44%) and mean age 24.79. Statistical analysis showed that this distribution was no significant difference ($P > 0.05$), [Table 4].

Regarding PCT there were significant differences ($P < 0.05$) observed between the mean serum concentration of PCT belong to 18 burn patients of 1st group (G1) (1.24 ± 0.82), 23 patients of 2nd group (G2) (1.52 ± 0.76), six patients of 3rd group (G3) (2.16 ± 0.11), and three burn patients of 4th group (G4) (6.5 ± 2.87) with mean serum concentration of 18 HC (0.02 ± 0.006). Significant differences obtained when compared to G3 (2.16 ± 0.11) versus G4 (6.5 ± 2.87) using F-test and *t*-test as shown in Tables 5 and 6.

DISCUSSION

Severe burn injury induces a distinct inflammatory response which is recognized by activation of all inflammatory pathways, dysregulation of cell-mediated immunity, and changes of mediators of the immune system including cytokines, vascular endothelium, growth factors, and different immunocompetent cell populations (Dehne et al., 2002). This study was carried out to investigate the role of PCT and other biological parameters as an inflammatory

Table 1: Gender distribution of burn patients

Gender	Burn patients	Chi-square <i>P</i> value
	No. (%)	
Female	32 (64)	0.14 $P>0.05$
Male	18 (36)	
Total	50 (100)	

$P>0.05$: No significant

Table 2: Burn patients groups according to the TBSA%

Whole groups	Total		Male		Female		Age (year)/Mean	TBSA%/Mean
	No	%	No	%	No	%		
G1	18	36	9	47.06	9	30.30	2.4–44 (18.08)	6–25 (16.67)
G2	23	46	7	41.18	16	48.48	1.2–39 (22.49)	26–50 (37.28)
G3	6	12	1	5.88	5	15.15	20–36 (28.17)	51–75 (59.17)
G4	3	6	1	5.88	2	6.06	16–38 (26.33)	76–100 (86.67)
Total	50	100	18	100	32	100	0–50 (22.23)	1–86 (35.03)

G1: (6–25%); G2: (26–50%); G3: (51–75%); G4: (76–100%)

TBSA: Total body surface area

Table 3: Mortality frequency and length of hospitalization of burned patients in both genders in relation to TBSA%

TBSA%	Gender		Total	Mortality		Total	Hospital stay (days) Mean±SE
	Female	Male		Female	Male		
G1	9	9	18	1	0	1	8.13 ± 2.97
G2	16	7	23	8	2	10	7.11 ± 3.13
G3	5	1	6	3	1	4	5.44 ± 4.41
G4	2	1	3	2	1	3	2.81 ± 2.15
Total	32	18	50	14	4	18	

G1: (6–25%); G2: (26–50%); G3: (51–75%); G4: (76–100%)

Female mortality 14 (28%); Male mortality 4 (8%)

Table 4: Distribution of survivor and non-survivor burn patients by gender

Gender	Survivors No.=32	Non-survivors No.=18	Chi-square	P value
	No. (%)	No. (%)		
Female	18 (56.25)	14 (77.78)	2.97	0.08
Male	14 (43.75)	4 (22.22)		
Total	32 (100)	18 (100)		

P>0.05: no Significant; Total mortality rate: of 18 (36%)

Table 5: Comparison estimation of mean serum concentration of PCT (ng/ml) in studied groups according to TBSA% using ANOVA test

Studied groups TBSA%	No.	Mean concentration of serum PCT (ng/ml)	P value F-test
		Mean±SE	
G1	18	1.24±0.82	0.016 S
G2	23	1.52±0.76	
G3	6	2.16±0.11	
G4	3	6.5±2.87	
HC	18	0.02±0.006	

S = Significant (P<0.05)

marker in burned patients. Regarding the distribution of burn trauma in relation to gender, this study showed no significant difference ($P > 0.05$) between male 18 (36%) and female 32 (64%) as shown in Table 1. In agreement with this finding most population-based studies in a developing country have found that the prevalence of thermal injury was higher in women than in men, this concept coincides with a previous study by Hosseini et al., 2007, who found that the most victim of burn injury were females and were caused by suicide attempt through the self-inflicted burn. Sadeghi-Bazargani et al. (2010) also reported that a

suicide attempt was common among females. However, the results of the present study were not compatible with those reported by Khan and Malik, 2006, Taghavi et al. (2010) who stated that the incidence of burn trauma among males was more than females. Indeed, the current study showed that females comprised 64% while male consist of 36% of admissions in WEEH in Erbil governorate, this high percentage of burn among female is due to family disturbances, which makes a large proportion of female to self-inflicted burn and major reason of marital conflict and improper education level. The highest number 16 of burn patients fell in 20–29 years of the age group, in which 13 (40.63%) in female and 3 (16.67%) in male, the mean age of burn patients was 22.2) years old. The finding of the present study was in agreement with those reported previously by other investigators Othman and Kendrick, 2010, Taghaddosinejad et al., 2010, who showed that the majority of burn patients were younger than 26 years old; they also reported that females were more exposed to burn agents in these age groups than in any other age groups. However, the results in this study contradict those reported by Rooh-ul-Muqim et al., 2007, who stated that children within 1–10 years of age group comprised the largest number of burn patients. The lowest percentage of burn patients, in the current study was fallen in age groups 40–49 years similar finding was reported by Maghsoudi et al., 2005 [Table 7]. Regarding the body extended to burn insult, the current study showed that mean of TBSA% burned was (35.03%) ranged (1–86%), in which the highest number 23 (46%) of burn victim fell in the 2nd group (G2) (26–50%), with mean age (22.49) years old and mean TBSA% burned (37.28%), while the lowest number of burn patients located in 4th group (G4)

3 (6%) with mean age (26.33) years old and mean TBSA% burned (86.67%), eventually another remaining number in current study among burn patients was 6 (12%) and 18 (36%) fell in 3rd group (G3) and 1st (G1) group, with mean age and mean TBSA% were (28.17), (59.17%), and (18.08), (16.67%) respectively [Table 2]. Similar finding was reported by several researchers Maghsoudi et al., 2005, and Jeschke et al., 2007, stated that out of 189 burn patients which were distributed into four groups depending on TBSA% burned, the highest number (79) burn victims fell in the second group with mean TBSA% (50%) ranged (<40–>80%). Throughout this study, the mean TBSA in admitted burnt is 35.03% ranged (6–100%). Although the majority of burn patients 23 (46%) had TBSA burnt (37.28%) located in 2nd group, there was a comparatively the small number of patients 3 (6%) with TBSA over 75%, all of whom died contributing to the high mortality in this research, this is mainly due to intentional self-harm burns with fire (flame burns). Out of 50 burn patient admissions, 18 patients died in the hospital giving a mortality rate of 36%. As shown in Table 3, seven of patients with a TBSA burnt over 50% in which three patients located in the 4th group with TBSA% (76–100%) burnt all of them died. The deceased number of burn patients was proportional to TBSA% burnt as depicted in Table 4. In addition, the length of hospital stay becomes lesser in patients with large burn size. The current study shows that mean length of stay

in various burn groups G1, G2, G3, and G4 was (8.13 ± 2.97), (7.11 ± 3.13), (5.44 ± 4.41), and (2.81 ± 2.15) days, respectively. Most studies from the EMR have reported that mean hospital stay was longer than this ranging from 11 to 16 days as reported by Maghsoudi et al., 2005, Jeschke et al., 2007. The reasons for the shorter hospital stay in this research may be due to higher rates of the mortality and inclusion of the larger number of self-harm burns than in other researches. As shown in Table 5, the highest frequent cause of burn was flame at age group (20–29) years, which denoted as the commonest age group. Finding of a recent study likewise revealed that scald burn comprised as a second commonest cause of the burn, followed by electrical and contact with hot object as an eventual cause of the burn. Similar results were reported by Hemeda et al., 2003, Carlos Siviero do Vale, 2005 who reported that flame was the most frequent cause of the burn.

CONCLUSION

PCT is a highly efficient laboratory parameter involving a simple and rapid bedside test for diagnosis and prognosis mortality in burn patients. The mean concentration of serum PCT shows significant differences when compared independently in deceased and survivor burn patients with HC.

CONFLICTS OF INTEREST

The author reports no conflicts of interest.

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Table 6: Comparison estimation of mean serum concentration of PCT (ng/ml) in studied groups according to TBSA % by using t-test

Studied groups TBSA%	P value/t-test	
G1 versus HC	0.221	NS
G2 versus HC	0.114	NS
G3 versus HC	0.130	NS
G4 versus HC	0.001	HS
G1 versus G2	0.777	NS
G1 versus G3	0.512	NS
G1 versus G4	0.006	HS
G2 versus G3	0.636	NS
G2 versus G4	0.008	HS
G3 versus G4	0.043	S

HS: Highly Significant ($P < 0.01$); S: Significant ($P < 0.05$); NS: Non-significant ($P > 0.05$)

Table 7: Distribution of burn patients in both genders in relation to age groups

Age group (year)	Burn patients No.50				Total	
	Female		Male			
	No.	%	No.	%	No.	%
1–9	3	9.37	3	16.67	6	12
10–19	8	25	7	38.89	15	30
20–29	13	40.63	3	16.67	16	32
30–39	6	18.75	4	22.22	10	20
40–49	2	6.25	1	5.55	3	6
Total	32	100	18	100	50	100

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