# Leukocytosis as Prognostic Factor for Severe Multiple Injuries

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## ABSTRACT:

**Objective:** The objective of this study was to determine if the white blood cell (WBC) count can predict severity of injuries and can be considered as a bad prognostic sign in multiple injured patients.

**Methods:** This was a comparative study comparing two groups of multiple injured patients according to the severity of injury, intensive care unit (ICU) admitted group with severe injuries and ward admitted one without significant injuries, and comparing their initial WBC in the first day. Also, the ICU admitted group was divided into two subgroups, survived and died one, comparing the decline in WBC count between them in the first 3 days.

**Results:** There was a difference in mean WBC count between ICU group with severe injuries and ward one without significant injury that was statistically significant (p<0.0001). Also there was a significant resistance of WBCs to decline to normal level in the died ICU subgroup as survived patients in the first three days (p<0.0001, p<0.0001 and p<0.0001 respectively).

**Conclusion:** A significant elevation in WBC count in severe multiple injured patients is found and it can be considered as a bad prognostic sign for those with slow decline to normal level within first three days.

**KEY WORDS**: multiple injured patient, leukocytosis, white blood cell count =WBC, ICU and ward.

## **INTRODUCTION:**

Emergency physicians are continually searching for early prognostic signs that efficiently differentiate trauma patients with major versus minor injury. Normal vital signs are not sufficiently sensitive to reliably exclude brain, bowel, vascular or solid organ injuries<sup>1-3</sup>. Lipsky showed that a significant proportion of trauma patients who eventually die had normal vital signs in the emergency department<sup>4</sup>.

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Theoretically, multiple injuries are considered as major stressful conditions, which induce neuronal and hormonal response, epinephrine and cortisol, producing leukocytosis from both bone marrow and splenic sources<sup>5-7</sup>. It has been hypothesized that elevation of trauma patient's WBC count may be a surrogate marker of neurohormonal activation and be valuable in identifying patients with injuries. Tissue major swelling (edema) occurring after trauma is probably an inflammatory response due to local cytokine production and increased leukocyte adhesion as a result of a direct effect on vascular permeability and leukocyte activation <sup>8-12</sup>. Since the complete blood analysis

is one of the first tests obtained from trauma patients in the emergency department, WBC count can serve as an easy obtained marker for a serious injury.Studies on blunt trauma patients have shown higher WBC counts in the more severely injured patients<sup>13-15</sup>. Although, Paladino et al showed that WBC count was not a useful addition as a diagnostic indicator of major trauma<sup>16</sup>, Rovlias et al in their study of head trauma patients in the neurosurgical intensive care unit (ICU) WBC showed that count was significantly higher in patients with severe head injury compared to those with minor to moderate injury<sup>17</sup>. In this study, we tried to find out whether there was a high WBC count associated with severity of injuries and can be considered a bad prognostic sign or not.

## **PATIENTS AND METHODS:**

The study was conducted at Alsadr Teaching Hospital, which receives approximately 100,000 yearly emergency-department visits and 5000 trauma activations per year. We measured the WBC counts of 334 (227 males and 107 females) severe and non-severe multiple injured patients, whom were admitted to the hospital between February 2009 and February 2013. Data collected included WBC count on admission, patient's age and (steroids, immunosex, drug suppressants, lithium, beta agonists) history, the Injury Severity Score (ISS) on arrival. Co-morbid conditions that might affect the degree of leukocytosis were recorded, including pregnancy, chronic infection. diabetes. cardiovascular disease. pulmonary disease, cancer, liver disease, pancytopenia on presentation, and immunologic diseases. Surgical procedures were excluded except emergency measures tracheostomy and chest tube. Also died patients at day arrival to hospital were excluded. The age range was 5 to 68 years and the mean age was 29.5.The WBC counts of the patients were obtained daily within the first three day and the patients were divided into two groups: First severe group who multiple injured patient admitted to ICU, 130 (96 male and 34 female) with either severe head injury Glasgow Coma Scale (GCS) scores less than 8 all intra-abdominal and intracranial injuries, spinal and skull fractures, pelvic diastasis, pulmonary contusions, hemothorax. pneumothorax, flail chest. The non-severe multiple injured group, 184 patient (131 male, 53 female) that admitted to the ward with minor injuries that do not meet the significant injury specifications outlined above, such as patients with head injury GCS above 8, nonemergent extremity fractures, as well as all patients, who did not develop significant injuries, had been admitted for observation.

### **RESULTS:**

A total of 334 blunt trauma patient records were reviewed. Of these, 130 were found to be severe multiple injured patients admitted to ICU and multiple iniured 184 non-severe patients admitted to ward. Of those patients with severe multiple injuries, fifty patients died in the ICU, 15 died in second day, 11 died in third day and 24 died after three days. Eighty severe multiple injured patients, whom were admitted to the ICU, survived, shifted to ward, and discharged home after observation. Of those with non-severe

multiple injuries, 184 were admitted for observation then discharged home. Table (1) shows a comparison between WBC count in severe and non-severe multiple injured patients. The mean WBC count in the severe multiple injured patients was  $17.94 \times 10^{9/1}$ compared to  $11.48 \times 10^9$ /l in the nonsevere multiple injured patients. The difference in WBC count values between the two groups was statistically significant (p<0.0001).

 Table (1): WBC count in severe (ICU group) and non-severe (ward group)

 multiple injured patients in the first day of admission

Severity	Ν	Mean	S.D	P value
ICU	130	17.94	3.565	0.0001
Ward	184	11.48	2.659	

The next step divided severe multiple injured group in to two subgroups, died and survived patients (Table 2). We compared WBC count for those two subgroups in the 1<sup>st</sup>, 2<sup>nd</sup> and 3<sup>rd</sup> days of admission. In the 1<sup>st</sup> day, the mean WBC count of died subgroup, 19.44 x  $10^{9}/1$ , compared to that of the survived subgroup, 17.00 x  $10^{9}/1$ . In the 2<sup>nd</sup> day, the mean WBC count of the died subgroup 16.00 x  $10^{9}/1$  compared to the

mean WBC count of the survived subgroup, 11.55 x  $10^9/1$ . After  $2^{nd}$  day, the mean WBC count of the died subgroup, 13.04 x  $10^9/1$  compared to the mean WBC count of survived subgroup 6.09 x  $10^9/1$ . There were statistically significant differences between WBC count for died subgroup as compared with survived subgroup in the first, second and after second days in the ICU (p<0.001, p<0.0001 and p<0.0001 respectively).

 Table (2): Comparison of WBC count in the survived to the died ICU patients in the first, second and after second days of admission

WBC counts	Fate	Ν	Mean	SD	P value	
Day 1	Survived	80	17.00	3.257	0.001	
	Died	50	19.44	3.552		
Day 2	Survived	80	11.55	2.690	0.0001	
	Died	35	16.00	3.134		
After Day 2	Survived	80	6.09	2.307	0.0001	
	Died	25	13.04	2.051		

## **DISCUSSION:**

Multiple injured patients seem to have elevated, or at least high normal, WBC count values on presentation. This suggests that the stress of trauma incident itself can result in marked demargination. The role of catecholamines and corticosteroids has reported been in the literature <sup>18</sup><sub>22</sub>.Catecholamines increase the leukocyte count by release of the marginated cells into the circulating pool. Corticosteroids increase the

neutrophil count by releasing the cells from the storage pool in the bone marrow into the blood and by

preventing egress from the circulation into these tissues  $^{18}2^{22}$ .

Another mechanism through which leukocytes number can be associated with tissue damage is the traumatic rupture of microvessels followed by physical occlusion. The leukocytes are less deformable than the erythrocytes, and a greater pressure gradient is therefore required to force them through the capillaries with small diameter. Under conditions of reduced perfusion pressure, the capillaries may be have liked a sieve and trap the leukocytes to increase the WBC number. After the entrapment, the leukocytes form a common area of contact with the endothelium and may not be dislodged even after the

perfusion pressure returns to normal <sup>23-</sup> <sup>26</sup>. The mechanical occlusion of the capillaries may become more evident as a result of the release of a number of cytotoxic chemicals that leads to increased leukocyte endothelial interactions <sup>27</sup>.In this study, blood WBC levels relationships are:

1- Patients with a mean WBC value of 17.94 x  $10^{9}$ /l were associated with severe multiple injuries, where as patients with a mean WBC value of 11.48 x  $10^{9}$ /l were associated with non-severe multiple injuries in first day of admission (Figure 1).

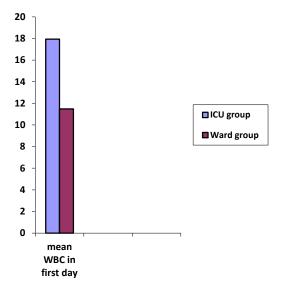
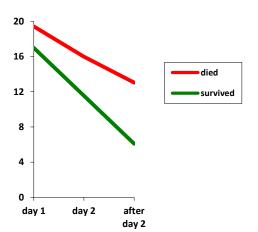


Figure (1): Diagram comparing mean WBC value of severe multiple injured patients to the mean WBC value of non-severe multiple injured patients in first day of admission.

2- Resistance of WBCs number to decline to normal level in the died subgroup. The mean WBC count 19.44 x  $10^{9}$ /l in the 1<sup>st</sup> day, 16.00 x  $10^{9}$ /l in the 2<sup>nd</sup> day and 13.04 x  $10^{9}$ /l in after 2<sup>nd</sup> day, can be considered as a bad prognostic sign for patients who died



in the ICU. Whereas, patients with rapid decline to normal WBC level in survived subgroup,  $17.00 \times 10^{9}$ /l in 1<sup>st</sup> day,  $11.55 \times 10^{9}$ /l in 2<sup>nd</sup> day and 6.09 x 10<sup>9</sup>/l in after 2<sup>nd</sup> day, was associated with a favorable outcome (Figure 2).

Figure (2): Diagram comparing the mean WBC count trend in died subgroup to the mean WBC count in the survived subgroup in the ICU.

Past studies studied several variables affecting WBC count, and have shown mixed results. Review of literature reveals several relevant studies:

 $al^{28}$ et performed Akköse а retrospective study evaluating 713 blunt trauma patients showing that count was proportionally WBC associated with ISS. Harris et al<sup>29</sup> retrospectively studied 46 patients after blunt abdominal trauma. They found that, in patients without obvious indications for invasive evaluation of the abdomen (e.g., peritoneal lavage, laparoscopy, laparotomy), leukocytosis was associated with intestinal injury. Paladino et al<sup>16</sup> in a heterogeneous blunt and penetrating trauma cohort (excluding isolated head injury) found that although there was a statistically significant higher WBC count in patients with major injuries, WBC

count was not a useful addition as a diagnostic indicator of major trauma in their study population. Gürkanlar et al<sup>30</sup> retrospectively studied 59 patients with head injury and found patients with severe head injury had significantly higher WBC count than those with moderate or minor injury.

#### **CONCLUSION:**

Elevated WBC count directly proportionally associated with severity of injury and those with slow decline of WBC count to the normal level carries a bad prognosis.

## **REFERANCES:**

**1**. Brown CV, Velmahos GC, Neville AL, et al. Hemodynamically "stable" patients with peritonitis after penetrating abdominal trauma: identifying those who are bleeding. Arch Surg. 2005; 140:767–72. [PubMed]

**2**. Adams SL, Greene JS. Absence of a tachycardic response to intraperitoneal hemorrhage. J Emerg Med. 1986;4:383–9. [PubMed]

**3**. Thompson D, Adams SL, Barrett J. Relative bradycardia in patients with isolated penetrating abdominal trauma and isolated extremity trauma. Ann Emerg Med. 1990;19:268–75. [PubMed]

**4**.Lipsky AM, Gausche-Hill M, Henneman PL, et al. Prehospital hypotension is a predictor of the need for an emergent, therapeutic operation in trauma patients with normal systolic blood pressure in the emergency department. J Trauma. 2006;61:1228–33. [PubMed]

**5**. Toft P, Helbo-Hansen HS, Tonnesen E, et al. Redistribution of granulocytes during adrenaline infusion and following administration of cortisol in healthy

volunteers. Acta Anaesthesiol Scand.1994;38:254–8. [PubMed]

**6**. Landmann R, Durig M, Gudat F, et al. Beta-adrenergic regulation of the blood lymphocyte phenotype distribution in normal subjects and splenectomized patients. Adv Exp Med Biol.1985;186:1051–62. [PubMed]

**7**.Bessey PQ, Watters JM, Aoki TT, et al. Combined hormonal infusion simulates the metabolic response to injury. Ann Surg. 1984;200:264–81. [PMC free article] [PubMed]

**8**. Dietrich WD, Chatzipanteli K, Vitarbo E, Wada K, Kinoshita K: The role of inflammatory processes in the pathophysiology and treatment of brain and spinal cord trauma. Acta Neurochir Suppl 89: 69-74, 2004.

**9**. Fee D, Crumbaugh A, Jacques T, Herdrich B, Sewell D, Auerbach D, Piaskowski S, Hart MN, Sandor M, Fabry Z: Activated/effector CD4+ T cells exacerbate acute damage in the central nervous system following traumatic injury. J Neuroimmunol 136: 54-66, 2003.

**10**. Gourin CG, Shackford SR.: Production of tumor necrosis factor-alpha and interleukin-l beta by human cerebral microvascular endothelium after percussive trauma. J Trauma 42:1101-1107,1997.

**11**. Juurlink BH: Introduction: The role of inflammation in mediating damage following stroke and neurotrauma. Brain Pathol 10: 93-94, 2000.

**12**. Lenzlinger PM, Hans VH, Joller-Jemelka HI, Trentz O, Morganti¬Kossmann MC, Kossmann T: Markers for cell-mediated immune response are elevated in cerebrospinal fluid and serum after severe traumatic brain injury in humans. J Neurotrauma 18: 479--489 30, 2001.

**13**. Morell V, Lundgren E, Gillott A. Predicting severity of trauma by admission white blood cell count, serum potassium level, and arterial pH. South Med J. 1993;86(6):658–9. [PubMed]

**14**. Santucci CA, Purcell TB, Mejia C. Leukocytosis as a predictor of severe injury in blunt trauma. West J Emerg Med. 2008;9:81–5. [PMC free article] [PubMed]

**15**. Holmes JF, Sokolove PE, Land C, et al. Identification of intraabdominal injuries in children hospitalized following blunt torso trauma. Acad Emerg Med. 1999;6:799–806. [PubMed]

**16**. Lorenzo Paladino, MD, Ramanand A. Subramanian, PhD, Elisabeth Bonilla, BS, and Richard H. Sinert, DO. Leukocytosis as Prognostic Indicator of Major Injury. Received December 10, 2009; Revised February 16, 2010; Accepted April 21, 2010.

**17**. Rovlias A, Kotsou S. The blood leukocyte count and its prognostic significance in severe head injury.Surg Neurol. 2001;55:190–6. [PubMed]

**18**. Boggs DR: The kinetics of neutrophilic leukocytes in health and in disease. Semins Hemat 4:359-386, 1967.

**19**. Capps JA: Astudy of the blood in general paralysis. Am J Med Sci 3:650-682, 1896 (cited in 1).

**20**. Clifton GL, Ziegler MG, Grossman RG: Circulating catecholamines and sympathetic activity after head injury. Neurosurgery 8:10-14,1981.

**21**. Czigner A, Mihaly A, Farkas O, Büki A, Krisztin-Peva B, Dobo E, Barzo P: Kinetics of the cellular immune response following closed head injury. Acta Neurochir (Wien) 149:281-289, 2007.

**22**. Dale DC: Leukocytosis, leukopenia, and eosinophilia. In: Harrison's, ed. Principles of Internal Medicine. New York: McGraw-Hill, Inc., 1991:359-362

**23**. Hallznbeck J, Dutka A, Tanishima T, Kochanek P, Kumaroo K, Thompson C, Obrenovitch T, Contrzras T: Polymorphonuclear leukocyte accumulation in brain regions with low blood flow during the early post-ischemic period. Stroke 17: 246-253, 1986

**24**. Janoff A, Schaeffer S, Scherer J. Bean MA: Mediators in inflammation in leukocyte lysosomes. Mechanism of action of lysosomal cationic protein upon vascular permeability in the rat. J Exp Med 132: 841-851, 1965.

**25**. Suval WD, Duran WN, Boric VIP, Hobson RV. Berendson PB. Ritter AB: Microvascular transport and endothelial cell alterations preceding skeletal muscle damage in ischernia and reperfusion injury. Am J Surg 154: 211-215, 1987.

**26**. Yamakawa T, Yamaguchi S, Niimi H. Sugiyama I: WBC plugging and blood flow maldistribution, in the capillary network of cat cerebral cortex in acute hemorrhagic hypotension: An intra-vital microscopic study. Circ Shock 22: 323-332, 1987.

27. Harlan JM: Leucocyte endothelial interactions. Blood 65: 513-525,1985.

**28**. Akköse S, Bulut M, Erol A, Veysel B, et al. Does the leukocyte count correlate with the severity of injury? Turkish Journal of Trauma and Emergency Surgery. 2003;9:111–113. [PubMed]

**29**. Harris HW, Morabito DJ, Mackersie RC, Halvorsen RA, et al. Leukocytosis and free fluid are important indicators of isolated intestinal injury after blunt trauma. J Trauma. 1999;46:656–659. [PubMed]

**30**. Gürkanlar, Predictive Value of Leukocytosis in Head Trauma. Turkish Neurosurgery 2009, Vol: 19, No: 3, 211-215

ارتفاع كريات الدم البيضاء عامل تنبؤي للإصاباتِ المتعدّدةِ الحادّةِ

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المحف: الهدف مـن هـذه الدراسـةِ هـي أَنْ تُقـرّرَ إذا ارتفـاع عـدد كريـة الـدمِّ البيضـاءِ يُمْكِنُ أَنْ يَتوقَـعَ شـدَّةَ الإصاباتِ ويُمْكِنُ أَنْ يُعتَبرَ كإشارة تنبؤية سيئة في المرضى المصابين المتعدّدين.

الطرق: هذا هي دراسة مقارنة ،تُقارنُ ارتفاع عدد كرية الدمِّ البيضاءِ في اليومِ الأولِ بين إثنان مِنْ مجموعاتِ المرضى المصابينِ المتعدّدينِ طبقاً لشدَّةِ الجرحِ، المجموعة الأولى تضم المرضى الراقدين في وحدة العنايةِ المركّزةِ والذين يعانون من اصابة شديدة مع المجموعة الثانية تضم المرضى الراقدين في الردهة والذين يعانون من اصابة غير شديدة.كما قسمت مجموعة المرضى الراقدين في وحدة العنايةِ المركّزة إلى مجموعتين فرعيتين، الناجين والمتوفين، قورن عدد كرية الدمِّ البيضاءِ بينهم في الأيام الأولى الثلاثة.

النَّتَائِج: كان هناك إختلاف هام في إحصاء عدد كرية الدمِّ البيضاء بين مجموعة وحدة العناية المركّزة والردهة للأصابات المتعددة الحادة (p <0.0001). أيضاً كان هناك a مقاومة هامّة عدد كرية الدمِّ البيضاء للإنجدار إلى مستوى طبيعي في الموتوفين مقارنة بالناجين من المرضى في العناية المركزة في الأيام الأولى الثلاثة (p 0.0001, p <0.001 على التوالي).

الحاتمة: الإرتفاع في عدد كرية الدمِّ البيضاءِ يُعتَبرَ إشارة تنبؤية سيئة للأصابات المتعددة الحادّةِ وخصوصا الذين ببطيئِ يَنحدرُ إلى مستوى طبيعي ضمن ثلاثة أيام أولى.