

Hematological profile of septicemia in elder age group patients

Delux V Godghate^{1*}, Vishal Narote², Arvind Bhake³

¹Assistant Professor, ²Junior Resident, ³Professor and Head, Department of Pathology, Jawaharlal Nehru Medical College, Sawangi (Meghe), Wardha, Maharashtra, INDIA.

Email: delux.v.g@gmail.com

Abstract

Background: Septicemia is a devastating medical condition encountered in many hospitals of developing countries. Septicemia in elderly patient is a critical issue due to high rates of morbidity and mortality. More than 60% of septicemia patients are over 65 years of age. The increasing incidence of sepsis among elderly individuals is a major public health concern. Therefore, early and accurate diagnosis is vital to ensure that appropriate medical decisions are made. To this end, clinicians in intensive care units and emergency departments need to predict systemic infection at an early stage. However, early diagnosis of sepsis is difficult because the signs and symptoms of sepsis in elderly patients, such as fever, are nonspecific and may be blunted or absent. Therefore, laboratory findings are important in the diagnosis of sepsis. **Aims and Objectives:** The present prospective observational study was undertaken to carry out simple, easy to perform hematological tests for detection of sepsis, to evaluate the validity of single test and to see the utility of C - reactive protein in septicemia, to find out the relationship between hematological parameters and bacteriologic ally proven sepsis. **Material and Methods:** Present study was done in the department of Pathology Central Laboratory, Jawaharlal Nehru Medical College and Acharya Vinoba Bhave Rural Hospital for a period of one year from December 2014 to December 2015. The study group consisted of 61 cases clinically suspected of septicemia in elders of the age 6 decades and above admitted in hospital. Two ml of Blood was collected in EDTA bulb from peripheral venipuncture and peripheral blood smear where made, stained with Leishman's stain and examined for any abnormality in morphology of blood cell. A complete blood count was performed by the cell counter (Backman coulter 5 AT diff). Laboratory investigation like CRP, micro ESR and blood culture were done. **Observation and Results:** After comparing the four test individually, latex -CRP showed sensitivity of (91.3.%) followed by Band /Neutrophil ratio (≥ 0.2) 78%, Leucopenia ($< 4000/\text{cu. mm}$) 63.1 % and Micro ESR (≥ 15 mm at one hour) 60.8% in identifying septicemia. Similarly more specificity was found in latex - CRP (80%) ,followed by Band /Neutrophil ratio (≥ 0.12) 67%, Leucopenia ($< 4000/\text{cu. mm}$) 66.6 % and Micro ESR (≥ 15 mm at one hour) 60% in identifying septicemia. **Conclusion:** Septicemia is a devastating medical condition encountered in many hospitals of developing countries where advanced diagnostic tools are not available in such situation hematological parameter like total leucocyte count, band / neutrophil ratio and morphologically, degenerative changes in neutrophil and serological test like CRP parameter can be easily obtained and used in early differentiation of septicemia from non septicemia cases in elderly patient as well help clinicians to diagnose septicemia. **Key Word:** septicemia, elder age.

* Address for Correspondence:

Dr. Delux V Godghate, Assistant Professor, Department of Pathology, Jawaharlal Nehru Medical College, Sawangi (Meghe), Wardha, Maharashtra, INDIA.

Email: delux.v.g@gmail.com

Received Date: 10/10/2016 Revised Date: 15/11/2016 Accepted Date: 18/12/2016

Access this article online

Quick Response Code:	Website: www.statperson.com
	DOI: 22 December 2016

INTRODUCTION

Septicemia in elderly patient is a critical issue due to high rates of morbidity and mortality. More than 60% of septicemia patients are over 65 years of age¹. The increasing incidence of sepsis among elderly individuals is a major public health concern. Therefore, early and accurate diagnosis is vital to ensure that appropriate medical decisions are made. To this end, clinicians in intensive care units and emergency departments need to predict systemic infection at an early stage. However,

early diagnosis of sepsis is difficult because the signs and symptoms of sepsis in elderly patients, such as fever, are nonspecific and may be blunted or absent^{2, 3}. Therefore, laboratory findings are important in the diagnosis of sepsis.

Laboratory parameter such as WBC count, differential count, C-reactive protein (CRP) level, procalcitonin level, and blood culture results can provide valuable information for the diagnosis of sepsis. WBC count is routinely recommended as an initial screening marker. Elevated WBC and absolute neutrophil count as well as left-shifted neutrophils are associated with sepsis⁴. These parameters can be determined using an automated hematology analyzer, but the morphology of neutrophils should also be reviewed through peripheral blood (PB) smears.

CRP has been shown to be a more sensitive and specific biomarker than WBC count⁵. Procalcitonin levels distinguish bacterial from viral infections, correlate well with clinical severity^{6, 7}, and show higher predictive sensitivity than CRP⁸. Blood culture remains the gold standard for diagnosing septicemia; however, this method shows low sensitivity⁹, requires an adequate amount of blood and proper collection techniques to avoid contamination, and can delay diagnosis as the results may require several days.

MATERIAL AND METHODS

Present study was done in the department of Pathology Central Laboratory, Jawaharlal Nehru Medical College and Acharya Vinoba Bhave Rural Hospital for a period of one year from December 2014 to December 2015. The study group consisted of 61 cases of elders of the age 6 decades and above admitted in hospital presenting with symptoms and signs of Fever, Mental confusion, Diarrhea Nausea, vomiting, Rapid pulse, Rapid breathing, low urine output, shock and clinical suspicion of septicemia admitted to department of medicine are included in the study group. The patient aged below 6 decades, any prior history of antibiotics therapy and patients who were not willing to participate were excluded. Detailed history and thorough examination was done in each case.

Two ml of Blood was collected in EDTA bulb from sample from peripheral venipuncture and peripheral blood smear where made, stained with Leishman's stain and examined for Red cells: morphology, immature forms, inclusion bodies, arrangement of cells, White cells: differential count, abnormal or immature forms like band cells and band/neutrophil ratio, micro ESR, Platelets for adequacy and abnormal forms and Parasites- malaria, filarial. A complete blood count was performed by the cell counter (Beckman coulter 5 AT diff).

Micro ESR was obtained by collecting capillary blood in a standard microhaematocrit tube (75mm length, internal diameter of 1.1mm and outer diameter of 1.5mm) and reading the fall of erythrocyte column after one hour. Value of more than 15mm was considered as suggestive of infection. CRP was done by latex agglutination slide test. The cut off value of 0.6 mg/dl as recommended by the manufacturer (Tulip diagnostics (p) Ltd, India) was considered as evidence of infection.

Blood culture was done in all the cases before starting antibiotics. Those having positive blood culture were taken as 'proved' sepsis and the remaining as 'probable' sepsis. Other relevant investigations like chest x-ray and lumbar puncture were done as and when required. Sensitivity, specificity, positive predictive value (PPV) and negative predictive value (NPV) of TLC, B/N ratio, ESR and CRP were determined to define the diagnostic significance of each test using cut-off values. Blood culture was used as the gold standard to consider a positive for sepsis.

OBSERVATION AND RESULTS

The present study comprised of 61 patients cases of elders of the age 6 decades and above admitted in hospital presenting clinically with symptoms and signs of septicemia. The age range of patient in this study was 60 to 80 years with a mean of 70 years. In this study out of 61 clinically suspected cases of septicemia 46 are proven to have septicemia and 15 unproved cases are placed under the category of probable septicemia as shown in Table -1. Out of 46 cases of proven septicemia having positive blood culture 11 died and out of 15 cases of probable septicemia 3 died as shown in Table -2. The Band/Neutrophil ratio of \geq to 0.12 was observed in 78 % of proven septicemia and 33.0% of probable septicemia cases. The overall positivity of the test was 67% in clinically suspected septicemia cases (Table-3). Similarly leucopenia ($<$ 4000/cumm) was found in 63.1% of proven septicemia and 33.3% of probable septicemia cases. The overall positivity for leucopenia was 55.7% in clinically suspected septicemia cases (Table-4).

The m-ESR was more than or equal to 15mm at one hour in 60.8 % of proven septicemia cases and 40% in clinically suspected septicemia cases. The overall positivity of the test was 55.7% in clinically suspected cases (Table-5). The latex agglutination test for CRP was positive (\geq to 0.6mg/dl) in 91.3% of proven septicemia and 20% of probable septicemia cases. The overall positivity in clinically suspected septicemia cases was 73.7% (Table-6). After comparing the four test individually, latex -CRP showed sensitive of (91.3%) followed by Band /Neutrophil ratio (\geq 0.2) 78%, Leucopenia ($<$ 4000/cu. mm) 63.1 % and Micro ESR (\geq

15 mm at one hour) 60.8% in identifying septicemia. Similarly more specificity was found in latex – CRP (80%) ,followed by Band /Neutrophil ratio (≥ 0.12) 67%,

Leucopenia ($<4000/\text{cu. mm}$) 66.6 % and Micro ESR (≥ 15 mm at one hour) 60% in identifying septicemia as shown in (Table 7).

Table 1: Distribution of patient

Clinically suspected septicaemia	61
Proven septicaemia	46
Probable septicaemia	15

Table 2: Blood culture and mortality in septicemia.

Blood culture	Positive	Negative	Total
No of patient	46 (75%)	15(25%)	61
Died	11(23.9%)	3(20%)	14

Table 3: Bands/Neutrophil count in septicemia

Band/ Neutrophil Ratio	Proven sepsis (n=46)		Probable sepsis (n=15)		Total no (n=61)		Control (n=61)	
	No	%	No	%	No	%	No	%
Band/ Neutrophil Ratio ≥ 0.12	36	78	5	33	41	67	2	3
Band/ Neutrophil Ratio < 0.12	10	22	10	67	20	33	59	97

Table 4: Leucocytes count profile in Septicemia

Leukocyte count (per cu mm)	Proven sepsis (n=46)		Probable sepsis (n=15)		Total no (n=61)		Control (n=61)	
	No	%	No	%	No	%	No	%
WBC <4000	29	63.1	5	33.3	34	55.7	6	9.8
WBC ≥ 4000	17	36.9	10	66.6	27	44.2	55	90.1

Table 5: Micro ESR Profile in septicemia

Micro ESR At one hour	Proven sepsis (n=46)		Probable sepsis (n=15)		Total no (n=61)		Control (n=61)	
	No	%	No	%	No	%	No	%
≥ 15 mm	28	60.8	6	40	34	55.7	5	8.1
< 15 mm	18	39.1	9	60	27	44.2	56	91.8

Table 6: CRP Profile (Latex Agglutination Method)

Latex CRP Test	Proven septicemia (n=46)		Probable septicemia (n=15)		Total no (n=61)		Control (n=61)	
	No	%	No	%	No	%	No	%
Positive (≥ 0.6 mg/dl)	42	91.3	03	20	45	73.7	2	3.2
Negative (< 0.6 mg/dl)	04	8.6	12	80	16	23.3	59	96.72

Table 7: Sensitivity and Specificity of four most useful individual tests

	Sensitivity (%)	Specificity (%)
Latex CRP (≥ 0.6 mg /dl)	91.3	80
Band /Neutrophil (≥ 0.12)	78	67
Micro ESR(≥ 15 mm at one hour)	60.8	60
Leucopenia ($<4000/\text{cu. mm}$)	63.1	66.6

DISCUSSION

Septicemia in elderly age patients remains a critical clinical issue because of the high rates of morbidity and

mortality. More than 60% of sepsis patients are over 65 years of age¹. In the present study out of 61 clinically suspected cases of septicemia 14 died. The definite diagnosis of septicemia is often difficult at early stage merely on clinical presentation due to non specific clinical presentation. Depending on signs and symptoms

clinicians can suspect septicemia but to establish diagnosis of septicemia laboratory test are essential. Various indirect markers of infection when studied collectively are very useful and reliable marker of septicemia.^{10,11} The predicative ability of this marker vary widely in different literature.¹²⁻¹⁵ Therefore, early and accurate diagnosis is vital to ensure that appropriate medical decisions are made. In our study we have done hematological investigations with special emphasis given to parameter like total leukocyte count, band /neutrophil ratio \geq to 0.12, degenerative changes in neutrophil (toxic granules, cytoplasmic vacuoles), m ESR \geq to 15 mm at first hour and serological investigation like C-Reactive protein \geq to 0.6 which can be done easily to see its usefulness in diagnosing the clinically suspected cases of septicemia. Sensitivity (diagnosing infection when it is present) and Specificity (not diagnosing infection when it is not present) of the test were calculated based on definition by Feinstein (1975) in detecting septicemia.

Elevated WBC and absolute neutrophil count as well as left-shifted neutrophils are associated with septicemia⁴ but present study showed leucopenia (55.7) with neutrophilia and increased in band/neutrophil ratio in 67% clinically suspected cases of septicemia. CRP test showed sensitivity 91.3% and specificity 80% in predicting septicemia. Similar observation were made in the study done by varsha, usha *et al*¹⁶. Leucopenia showed sensitivity 63.1% and specificity 66.6% in predicting septicemia while sensitivity of 78% and specificity of 67% was observed in band/neutrophil ratio. Study done by Philip AGS, Hewitt JR *et al* correlated leucopenia and increased band neutrophil ratio in bacterial infection¹³.

Hence in the present study we found that in a clinically suspected cases of septicemia hematological parameter like total leucocytes count, band / neutrophil ratio and morphologically, degenerative changes in neutrophil and serological test like CRP which can be done easily are helpful for clinician in early diagnosis of septicemia.

This study is a prospective observational study and is limited by its small sample size and highly selected group. These limitations may have led to incorporation bias overestimating the diagnostic power of the investigated markers. Larger prospective cohort studies are needed to further validate the clinical usefulness of these parameters.

CONCLUSION

Septicemia is a devastating medical condition encountered in many hospitals of developing countries where advanced diagnostic tools are not available in such situation hematological parameter like total leucocytes

count, band / neutrophil ratio and morphologically, degenerative changes in neutrophil and serological test like CRP parameter can be easily obtained and used in early differentiation of septicemia from non septicemia cases in elderly patient as well help clinicians to diagnose septicemia.

REFERENCES

1. Angus DC, Linde-Zwirble WT, Lidicker J, Clermont G, Carcillo J, Pinsky MR. Epidemiology of severe sepsis in the United States: analysis of incidence, outcome, and associated costs of care. *Crit Care Med*. 2001;29:1303–1310.
2. Gleckman R, Hibert D. Afebrile bacteremia. A phenomenon in geriatric patients. *JAMA*. 1982;248:1478–1481.
3. Castle SC, Norman DC, Yeh M, Miller D, Yoshikawa TT. Fever response in elderly nursing home residents: are the older truly colder? *J Am Geriatric Soc*. 1991;39:853–857.
4. Wile MJ, Homer LD, Gaehler S, Phillips S, Millan J. Manual differential cell counts help predict bacterial infection. A multivariate analysis. *Am J Clin Pathol*. 2001;115:644–649.
5. Isaacman DJ, Burke BL. Utility of the serum C-reactive protein for detection of occult bacterial infection in children. *Arch Pediatr Adolesc Med*. 2002;156:905–909.
6. Fernandez Lopez A, Luaces Cubells C, Garcia Garcia JJ, Fernandez Pou J. Procalcitonin in pediatric emergency departments for the early diagnosis of invasive bacterial infections in febrile infants: results of a multicenter study and utility of a rapid qualitative test for this marker. *Pediatric Infect Dis J*. 2003;22:895–903.
7. Kourtis AP, Sullivan DT, Sathian U. Practice guidelines for the management of febrile infants less than 90 days of age at the ambulatory network of a large pediatric health care system in the United States: summary of new evidence. *Clin Pediatr (Phila)* 2004;43:11–16.
8. Yo CH, Hsieh PS, Lee SH, et al. Comparison of the test characteristics of procalcitonin to C-reactive protein and leukocytosis for the detection of serious bacterial infections in children presenting with fever without source: a systematic review and meta-analysis. *Ann Emerg Med*. 2012;60:591–600.
9. Procop GW, Hartman JS, Sedor F. Laboratory tests in evaluation of acute febrile illness in pediatric emergency room patients. *Am J Clin Pathol*. 1997;107:114–121.
10. Da Silva O, Ohlsson A, Kenyon C. Accuracy of leukocyte indices and CRP for diagnosis of neonatal sepsis: a critical review. *Pediatric Infect Dis J* 1995; 14:362-6.
11. Kuruvilla KA, Pillai S, Jesudason M, Jana AK. Bacterial profile of sepsis in a neonatal unit in south India. *Indian Pediatr* 1998; 35:851-8.
12. Manroe BL, Weinberg AG, Rosenfeld CR, Browne R. The neonatal blood count in health and disease. 1. Reference values for neutrophilic cells. *J Pediatr* 1979; 75:89-98.
13. Philip AGS, Hewitt JR. Early diagnosis of neonatal sepsis. *Pediatrics* 1980; 65: 1036- 41.

14. Liu HC, Lehan C, Speer ME, Fernbach DJ, Arnold J. Rudolph AJ. Degenerative Changes in neutrophils: an indicator of bacterial infection. *Pediatrics* 1984; 74:823-827.
15. Mehboob, R. *et al* Deranged biochemical and hematological profile of septicemia patients in Mayo hospital, Lahore. 2016. Punjab Univ. *J.Zool.*, **31**(1): 87-93.
16. Varsha, Usha Rusia, Meera Sikka et.al, Validity of hematological parameters in identification of early and late onset neonatal infection. *Ind.J.Pathol.Microbiol.* 2003;46:565-568.

Source of Support: None Declared
Conflict of Interest: None Declared