# OUTCOME OF SEPSIS IN POST OPERATIVE ADULTPATIENTS OF CARDIAC SURGICAL INTENSIVE CARE UNIT

#### Rehana Javaid, Syed Muzaffar Hassan, Imran Bashir, Safdar Ali Khan, Irie Pervez, Safdar Abbas, Maha Bhatti, Maham Zahid, Hafsa Khalil, Razia Javaid

Armed Forces Institute of Cardiology/National Institute of Heart Diseases/National University of Medical Sciences (NUMS) Rawalpindi

Pakistan

## ABSTRACT

**Objective:**To determine the frequency of sepsis and to evaluate the morbidity and mortality of patients subjected to cardiac surgery that developed sepsis during the in-hospital postoperative period.

Study Design: Descriptive cross sectional study.

*Place and Duration of Study:* Armed Forces Institute of Cardiology, Intensive Care Unit of Rawalpindi, from Oct2016 to Dec2016.

*Material and Methods:*Three hundred and twenty eight patients were included who underwent elective open heart surgery at intensive care unit of AFIC/NIHD. All the patients who were above 18 years of age and underwent elective coronary artery bypass graft (CABG) surgery during the specified study period were included. Patients with any infection and with infective endocarditis were excluded. Non-probability consecutive sampling technique was used.

**Results:** The frequency of immediate post surgical sepsis was found to be 55(17.0%). The mean age of patients was found to be  $55(53\pm14)$  years. Among these patients, 29(9.0%) were males and 26(8.0%) were females. 38 (12.0%) had coronary artery bypass graft (CABG) and 17 (5.0%) underwent valve replacement. Mean ICU stay was found to be  $14.5 \pm 10.1$  days in males &  $17.3 \pm 12.4$  days in females, surgical site infection rate 8 (2.3%), cardiac support 19 (34.5%), likewise mechanical cardiac support in males 10 (18.0%) and in females 2 (4.0%), mean mechanical ventilation timewas found to be  $(5.1\pm1.3 \text{ hours})$ , post-op acute renal failure in males 6 (11.0%) and 1(2.0%) in females. Mortalityin males was found to be 5 (9.0%)and in females it was 6(11.0%). Haemocultures results of septic population showed that 15 patients were positive and of which 11 (73.0%) mainly comprised of the acinetobacter species.

**Conclusion:**Sepsis that occurs in the cardiac surgery post-operative period significantly contributes to patients' high morbidity and mortality. Moreover, further studies are essential to simplify the proper diagnosis, and risk factors that will help prevent sepsis.

Keywords: Cardiac surgery, Postoperative period, Sepsis.

This is an Open Access article distributed under the terms of the Creative Commons Attribution License (http://creativecommons.org/licenses/by/4.0), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

### INTRODUCTION

physiologic, Sepsis. syndrome of а pathologic, and bio-chemical abnormalities caused by infection, is a main public health concern, accounting for more than \$20 billion (5.2%) of total US hospital costs in 2011<sup>1,2</sup>. Sepsis is linked with high rates of morbidity and mortality, responsible for as much as one of every two to three in-hospital deaths. Particularly, the mortality rates of sepsis amplified during the last

**Correspondence: Dr Rehana Javaid**, Armed Forces Institute of Cardiology/NIHD RawalpindiPakistan *Email: dr.rehanajavaid@yahoo.com* 

decade, which is in contrast to the declining rates of all other major causes of death in the US<sup>2</sup>. Sepsis after cardiac surgery has been prescribed as a low-prevalence infectious complication with terribleresults. Patients who develop sepsis, regardless of the infectious focus present with high morbidity and mortality, which show a discrepancy from 17% to 65%<sup>3</sup>. Finding the incidence of sepsis is of great concern to both clinicians and public health officials, in order to calculate the load of the disease<sup>4</sup>. However, assessment of sepsis incidence is difficult, as it depends on the definition of sepsis, the method used to evaluate the condition, and the underlying population. Until 1992, no consensus existed on the terminology used to describe the presence and severity of sepsis, impairing comparison of studies on sepsis incidence and therapy outcomes<sup>5</sup>. This issue was addressed by The College American of Chest Physicians/Society of Critical Care Medicine (ACCP/SCCM) Consensus Conference in 1991, with the purpose to create a set of criteria for identifying and assessing the severity of sepsis<sup>6</sup>. The consensus proposal incorporated a introduction of the systemic inflammatory response syndrome (SIRS) criteria for early

pulmonary specialists in January 2014<sup>15</sup>. In 2016 the definition of sepsis was rationalized to categorise sepsis as a life-threatening organ dysfunction caused by a dysregulated host response to infection (by The Third International Consensus Definitions for Sepsis and Septic Shock)<sup>16</sup>.

Body mass index  $\geq 40 \text{kg/m}^2$ , haemodialysis in the preoperative period, pre-op cardiogenic shock, age ≥80 years, pre-op treatment with immunosuppressive agents, diabetes mellitus, mechanical circulatory support, 3 or more

\_

Table-I: Demographics and basic clinical information.	
Variables	Sei

Variables	Sepsis Cases
Gender, [n (%)]	
Male=245 (72.0%)	55 (17.0%)
Female=95 (28.0%)	
Hypertension, n (%)	
153 (45.0%)	30 (9.0%)
Diabetes mellitus, n (%)	
92 (27.0%)	19 (6.0%)
Type of surgery, n (%)	
CABG = 246 (72.0%)	38 (12.0%)
Valve = 60 (18.0%)	17 (5.0%)

identification of sepsis, defining sepsis as 2 SIRS criteria in patients with known or suspected infection, and severe sepsis as sepsis associated with organ dysfunction, hypoperfusion, or hypotension. However, repeatedly criticised for being too sensitive<sup>7,8</sup> and of questionable prognostic value<sup>9,10</sup> these easily applied "Bone criteria" remained the clinical standard in many hospital guidelines even after the introduction of internationally agreed-upon, but more comprehensive, criteria<sup>11-13</sup>. A 2001 task force, identifyingboundaries with these definitions, extended the list of diagnostic criteria but did not suggest alternatives because of the lack of underneath evidence<sup>14</sup>.and PIRO criteria was introduced. It was Recognized to reexamine the current definitions, the European Society of Intensive Care Medicine and the Society of Critical Care Medicine convened a task force of 19 critical care, infectious disease, surgical, and

revascularized vessels are the most important predictors of infections in postoperative period<sup>17</sup>.

Our study was conducted according to latest definition by the Third International Consensus Definitions for Sepsis and Septic Shock 2016, in which sepsis is defined as "a life-threatening organ dysfunction caused by a dysregulated host response to infection".

For clinical investigation, organ dysfunction can be represented by an increase in the Sequential (Sepsis-related) Organ Failure Assessment (SOFA) score of 2 points or more, which is associated with an in-hospital mortality greater than 10%.

# MATERIAL AND METHODS

From October 2016 to December 2016, 328adult patients who underwent cardiac surgery were included in this study. Prophylactic antibiotic therapy was given according to the

protocol of our hospital, from the induction of anesthesiaup to ICU stay. Selection of patients was done by Armed Forces Nursing staff that were trained and informed about the inclusion and exclusion criteria. Sepsis in the postoperative period was defined as confirmation of infection linked with two or more criteria of systemic inflammatory response syndrome: bodv temperature >38°C or <36°C, leukocytes >12,000 cells/mm<sup>3</sup>, positive blood cultures, and respiratory rate >20/min, heart rate >100/min. According to new definition of sepsis, Organ spent in the ICU, surgical site infection rate, cardiac support, mechanical cardiac support,mechanical ventilation time, positive haemocultures, post-op acute renal failure and mortality.

## RESULTS

There were 328 cardiac surgery patients subjected to cardiac surgery during the studyperiod, 246(75%) had CABG and 62(18%) underwent valve surgeries.Among these patients,245 (72.0%) were males and 95 (28.0%) were females. Demographics of patients are

Table-II: Septic patients in cardio surgical icu with demographics and clinical findings.

Variables	Males n=29 (9.0%)	Females n=25 (8.0%)		
Age (mean±SD)	55 (51±15)	60 (58±8)		
ICU stay(mean )	14.5 ±10.1 days	17.3 ± 12.4 days		
Cardiac supportn(%)	10(18.0%)	09 (16.0%)		
Mechanical cardiac	10 (18.0%)	2(4.0%)		
supportn(%)				
Mechanical ventilation	5 hours	5 hours		
time (median)				
CRRT n(%)	6(11.0%)	1(2.0%)		
Surgical site infection rate	6(11.0%)	2(4.0%)		
(SSI) n(%)				
Mortality n(%)	05(9.0%)	6(11.0%)		
Table-III: Isolated microorganisms from haemocultures of septic patients.				
Isolated Microorganism	Number	Number of patients with positive haemoculture		
	(expressed as	a percentage of total septic patients) (n=15)		
Acinetobacter spp.		11 (73.0%)		
Klebsiella pneumoniae		02 (13.0%)		
E.coli		01 (7.0%)		
Burkhuldoria cepacia		01 (7.0%)		

dysfunction can be recognized as an acute change in total SOFA score>2 points consequent to the infection. The baseline SOFA score are supposed to be zero in patients not known to have preexisting organ dysfunction. ASOFA score<2 reflects an overall mortality riskof about 10% in a general hospital population with assumed infection. In our study, after diagnosis of sepsis, identified septic patients were analysed usingdescriptive statistical tests i.e.chi square and t-test. The assessed characteristics of the patients with sepsis were: age, gender, number of days shown in table-I.

Fifty five(17%) developed sepsis during the in-hospital post-op period. There was a predominance of male patients with sepsis during the post-op period when compared to female patients (69% vs. 31%); the patients' mean age was 55 ( $53\pm13$ ) years. Mean ICU stay was found to be 14.5 ± 10.1 days in males & 17.3 ± 12.4 days in females, surgical site infection rate 8 (2.3%), cardiac support 19 (34.5%), likewise mechanical cardiac support in males 10 (18.0%) and in females 2 (4.0%), mean mechanical

ventilation time was found to be  $(5.1\pm1.3 \text{ hours})$ , post-op acute renal failure in males 6 (11.0%) and 1(2.0%) in females. Mortality in males was found to be 5 (9.0%) and in females it was 6(11.0%). Haemocultures results of septic population showed that 15 patients were positive and of which 11 (73.0%) mainly comprised of the acinetobacter species (table-II).

Isolated microorganisms from haemocultures of septic patients are shown in table-III. All of them received antibiotic therapy according to antibiogram. Of all 55 septic patients 11 (3.3%) of them died during ICU stay. No Methicillin Resistant Staphylococcus Aureus (MRSA) was found.

# DISCUSSION

The prevalence of infectious complications after cardiac surgery is up to 3.5% of patients. Cardiac surgery postoperative infections increase the morbidity and mortality, hospital cost and ICU stay duration<sup>18</sup> and it is similar with finding of our study i.e.frequency of sepsis in our study was 55 (17.0%).

Cardiac dysfunction is an important component of multiorgan failure that is caused by severe sepsis<sup>19,20</sup>. Regarding multi organ failure, continuous renal replacement therapy (CRRT) was performed on 7 (2%) patients.Patients with known organ failure disease who then become septic appear to have increased mortality. Overall mortalitycalculated in our study was 24 (7%). In which 11 (20.0%) were from sepsis group<sup>20</sup>. These findings argue for the critical importance of early identification and aggressive response to sepsis in cardiac patients, given that they appear to be at increased risk for complications. In another study, the sepsis associated mortality is 27% but exceeds 50% in cases of septic shock<sup>21,22</sup>.

Frequency of sepsis in our study was 55 (17.0%), which is associated with hospital cost and occurrence of sepsis . Toumpoulis et al<sup>23</sup>. Studied 3,720 patients submitted to cardiac surgery, with the objective of identifying risk factors for sepsis and endocarditis. The

prevalence of sepsis in the post-op period was 1.2%; however, the in-hospital mortality was >70% and there was an increase in cost and time of hospitalization associated with the pace of sepsis<sup>23</sup>.

The Surviving Sepsis guidelines additionally stress the importance of initiating antibiotics within an hour of identification of septic shock, with empiric therapy targeted to the suspected site of infection and responsible organisms, and then working to obtain infection source control as quickly as possible<sup>24</sup>. Overall surgical site infection rate during study time period was found to be 8 (2.3%). Out of this, 6 (62%) patients were from sepsis group and it was in accordance with the present study.

Results of a study reported by Stoeckle etal<sup>25</sup>, conclude that relative frequency of bloodstream infections is found be more than 4 times higher in diabetic than in non-diabetic general inpatients. Diabetic population of our study was19 (35.0%). This finding was similar with previous literature.

According to the third international consensus definitions for sepsis and septic shock, the sequential onset of complications, that is, the occurrence of adverse events after the onset of sepsis, proposed that the patients with this disease are more vulnerable and thus present such high rates of complications. This lends to belief that the patient with sepsis has a higher chance of presenting with acute renal failure, heart failure. An added challenge is that infection is seldom confirmed microbiologically when treatment is started; even when micro-biological tests are completed, culture-positive "sepsis" is observed in only 30% to 40% of cases<sup>16</sup>. This is similar in the present study, only 15 microorganisms were isolated which are mentioned in table-III.

In spite of its low prevalence, sepsis that occurs in the cardiac surgery post-operative period significantly contributes to patients' high morbidity and mortality. Therefore, the current recommendations for the treatment of severe sepsis represent a strategy that is potentially capable of improving the outcome of such patients and needs to be evaluated concerning the sepsis that occurs after cardiac surgery.

### CONCLUSION

Sepsis has always been a rare complication in the postoperative period of cardiac surgery. However, when observed, the result is catastrophic: 7% of the patients who presented such complication died during the hospitalization phase. Cardiac surgery, in most cases, is a clean surgery with a low rate of infectious complications; however, when these do happen, they contribute to patients' unfavorable outcome. The therapeutic measures suggested by the international guidelines for the treatment of sepsis must be functional to patients that present with sepsis after cardiac surgery, as they are potentially proficient of reducing morbidity and mortality.

#### Limitation of the Study

The study was conducted at single center with relatively small sample size. A multicenter study with larger sample size is needed.

## **CONFLICT OF INTEREST**

This study has no conflict of interest to declare by any author.

#### REFERENCES

- 1. Liu V, Escobar GJ, Greene JD. Hospital deaths in patients with sepsis from 2 independent cohorts. JAMA 2014;312: 90–2.
- Wang HE, Devereaux RS, Yealy DM, Safford MM, Howard G. National variation in United States sepsis mortality: a descriptive study. Int J Health Geogr 2010;9:9.
- Silva E, Fernandes Jr CJ, Akamine N, Sogayar AMCB, Knobel E. Sepse e choque séptico. In: Knobel E. Condutas no paciente grave. 3ª ed. São Paulo: Editora Atheneu 2006. p. 61-78.
- Dellinger RP, Levy MM, Rhodes A, Annane D, Gerlach H, Opal SM, et al. Surviving sepsis campaign: international guidelines for management of severe sepsis and septic shock, 2012. Intensive Care Med 2013;39:165–228.
- Bone RC. Sepsis, the sepsis syndrome, multi-organ failure: a plea for comparable definitions. Ann Intern Med 1991;114:332–3.

- Bone RC, Balk RA, Cerra FB, Dellinger RP, Fein AM, Knaus WA, et al. Definitions for sepsis and organ failure and guidelines for the use of innovative therapies in sepsis. The ACCP/SCCM consensus conference committee. American college of chest physicians/society of critical care medicine. Chest 1992; 101: 1644–55.
- Levy MM, Fink MP, Marshall JC, Abraham E, Angus D, Cook D, et al. 2001 SCCM/ESICM/ACCP/ATS/SIS International Sepsis Definitions Conference. Intensive Care Med 2003;29: 530–8.
- Dear VJL, SIRS I'm sorry to say that I don't like you. Crit Care Med 1997;25:372-4.
- Pittet D, Rangel-Frausto S, Li N, Tarara D, Costigan M, Rempe L, et al. Systemic inflammatory response syndrome, sepsis, severe sepsis and septic shock: incidence, morbidities and outcomes in surgical ICU patients. Intensive Care Med 1995;21: 302–9.
- Salvo I, de Cian W, Musicco M, Langer M, Piadena R, Wolfler A, et al. The Italian SEPSIS study: preliminary results on the incidence and evolution of SIRS, sepsis, severe sepsis and septic shock. Intensive Care Med 1995;21(Suppl 2): S244–9.
- Shapiro N, Howell MD, Bates DW, Angus DC, Ngo L, Talmor D. The association of sepsis syndrome and organ dysfunction with mortality in emergency department patients with suspected infection. Ann Emerg Med 2006;48:583–90.
- Mayo Clinic Staff. Sepsis Symptoms Mayo Clinic [Internet]. 2014 [cited 2015 Sep].
- 13. UCLA Medical Center. Sepsis\_Nurse\_Driven\_Protocol [Internet]. 2013 [cited 2015 Sep].
- LevyMM, FinkMP, Marshall JC. International Sepsis Definitions Conference. 2001 SCCM/ESICM/ACCP/ATS/SIS International Sepsis Definitions Conference. Intensive CareMed 2003; 29(4):530-538.
- Vincent J-L, Opal SM, Marshall JC, Tracey KJ.Sepsis definitions: time for change. Lancet 2013;381(9868):774-775.
- 16. Singer M, Deutschman CS, Seymour C. The third international consensus definitions for sepsis and septic shock (sepsis-3) JAMA 2016;315:801–10.
- Fowler VG, O'Brain SJ, Muhlbaier LH, Corey GH, Ferguson BT, Peterson ER. Clinical predictors of major infections after cardiac surgery. Circulation 2005;(1):112: 1358-1365.
- Slaughter MS, Olson MM, Lee JT Jr, Ward HB. A fifteen years wound surveillance study after coronary artery bypass. Ann Thorac Surg1993;56(5):1063-1068.
- Court O, Kumar A, Parrillo JE. Clinical review: myocardial depression in sepsis and septic shock. Crit Care 2002;6:500–8.
- Zanotti-Cavazzoni SL, Hollenberg SM. Cardiac dysfunction in severe sepsis and septic shock. Curr Opin Crit Care 2009;15:392–7.
- Toumpoulis IK, Anagnostopoulus CE, Toumpuolis SK, De Rose Jr JJ, Swistel DG. Risk factors for sepsis and endocarditis and long-term survival following coronary artery bypass grafting. World J Surg 2005;29(5):621-627.
- Dellinger R.P. Surviving sepsis campaign: international guidelines for management of severe sepsis and septic shock: 2012. Crit Care Med 2013. 41(2): p. 580-637.
- Vincent JL, Sakr Y, Sprung CL, Ranieri VM, Reinhart K, Gerlach H, et al. Sepsis in European intensive care units: results of the SOAP study. Crit Care Med 2006;34:344–53.
- Brun-Buisson C, Doyon F, Carlet J, Dellamonica P, Gouin F, Lepoutre A, et al. Incidence, risk factors, and outcome of severe sepsis and septic shock in adults. A multicenter prospective study in intensive care units. French ICU Group for Severe Sepsis. JAMA 1995;274:968–74.
- Stoeckle M, Kaech C, Trampus A, Zimmerli W. The role of diabetes mellitus in patients with blodstream infections. Swiss Med Wkly 2008;138:512–9.

.....