Original Article

A study of echocardioghraphic indices in patients of sepsis and severe sepsis

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Abstract

Introduction: Patients with sepsis have a markedly depressed LV ejection fraction, a low cardiac output as measured by the Doppler technique, and systemic vascular resistance higher than usual in sepsis. Aims and Objectives: To evaluate the various echocardiographic indices in patients of sepsis, severe sepsis and septic shock. To correlate these various clinical parameters and echocardiographic indices with the outcome. Material and Methods: This hospital based prospective analytic observational study was performed in the parent institute from June 2009 to November 2010. A total of 70 cases of sepsis who were admitted in Medical ward or in the medical ICU were studied. INCLUSION CRITERIA: All patients satisfying the criteria for sepsis admitted to ICU and Medicine Wards. Observation and Results: This was a hospital based observational study which was done with 70 patients of sepsis who were admitted in Medicine wards or in the Medical ICU of the parent Institute. Out of 70 cases studied 50 were males as compared to 31 females. Majority of cases were between age group of 41-50 years. Mean age of the cases was 43.2± 15.3 years. The most common symptom was fever which was present in 85% of the patients,. Of the 70 subjects included in the study, 26(37.1%) had presented with sepsis, 32(45.7%) had severe sepsis and 12 (17.2%) patients were in septic shock. 46 (66%) survived and 24 (34%) succumbed to the illness. The most common etiology in severe sepsis was consolidation and was present in 47% of the patients.. In septic shock, the most common etiology and clinical presentation was consolidation (58%). In the non survivors, admission EF \geq 40 were found in 21 patients (87%) and, EF \leq 40 were found in 3 patients (13%). In the survivors, EF \ge 40 were found in 29 patients (63%) and, EF \le 40 were found in 17 patients (37%). Thus in our study 17 patients died with Distributive Shock (71%) and 7 patients (29%) died with Cardiogenic Shock had myocardial depression. Conclusion: The mortality was more in the septic shock patients (83%) as compared to those with severe sepsis (43.7%). Ejection Fraction and Cardiac Output was higher in Non survivors than Survivors at time of admission. However in non survivors, there were progressive decline, in ejection fraction over a period of stay but still their cardiac output was in normal range.

Keywords: echocardioghraphic, sepsis.

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INTRODUCTION

Severe sepsis is characterised by concomitant organ dysfunction and septic shock results when blood pressures fall despite adequate fluid resuscitation. Severe sepsis or septic shock accounts for as many deaths as

acute myocardial infarction in hospitals.2 The typical hemodynamic profile in early sepsis is the peripheral vasodilatation, systolic alterations of left ventricular and right ventricular which along with increased vessel permeability leads to hypovolemia and hypotension. Patients with sepsis have a markedly depressed LV ejection fraction, a low cardiac output as measured by the Doppler technique, and systemic vascular resistance higher than usual in sepsis. nowadays it is advised that all patients hospitalized with septic shock should be monitored using echocardiography as a noninvasive bedside procedure. As no Indian studies echocardiography in sepsis were found, this study was undertaken to evaluate the cardiac functions in patients of sepsis in our setup.

MATERIAL AND METHODS

This hospital based prospective analytic observational study was performed in the parent institute from June 2009 to November 2010. A total of 70 cases of sepsis who were admitted in Medical ward or in the medical ICU were studied.

METHODOLOGY

- 1. Study Design: Hospital Based Prospective Analytic Observational Study.
- 2. Number of study subjects: 70
- 3. Inclusion Criteria: All patients satisfying the criteria for sepsis admitted to ICU and Medicine Wards.
- 4. Exclusion Criteria: Following patients were excluded from the study.
 - a. Patients with known cardiac disease like rheumatic heart disease, ischemic heart disease, cardiomyopathy.
 - b. Patients who were having ECG changes of ischemic heart disease.
 - c. Patients who were on ventilatory support were also excluded from the study.

Statistical Analysis

Statistical analysis was performed with the help of SPSS 13.0 software. Statistical analysis included the usual

descriptive and univariate analysis. Student 't' test was used to compare continuous variables. For categorical variables, Chi-square,T test was used and unadjusted odds ratio with 95% confidence interval (CI) were calculated and 'p' values were calculated. In addition to the usual descriptive and univariate analysis, multivariate analysis was also performed. Multiple logistic regression analysis was done to find out association of variousclinical parameters with outcome of sepsis.

OBSERVATIONS AND RESULTS

Table 1: Age and sex distribution of the Sepsis patients

Sr. no.	Age in yrs	Male	Female	Total
1	<20	3	1	4
2	21-30	12	3	15
3	31-40	6	6	12
4	41-50	8	8	16
5	51-60	13	1	14
6	>60	8	1	9
Total		50	20	70

Maximum number of patients were in the age group of 41-50 years. The mean age of the study subjects was 43.2+15.3 years. There were 50M and 20F in the study.

 Table 2: Serial echocardiographic measurements of in terms of survival and non-survival

		SURVIVERS			NON SURVIVERS			
		46			24			
		DAV 1	DAY3	DAY 5	DAY 1	DAY 3	DAY 5	
		DAY 1	DATS		24	19	7	
	Lvidd	46.5± 7.1	44.3±3.8	44.8±3.4	47.1±6.5	48.1±4.3	49.8±4.4	
	Lvids	33.2±7.7	32±4.9	32.9±3.9	31.6±5.5	34.3±3.9	38.8±6.2	
Ejection	fraction (%)	50±14	50±13	49±9	57±12	48±11	42±17	
Strok	e volume	60.2±19.9	52.5±13.51	55±10.4	74.2±27.9	68.4±22.01	58.6 ±14.7	
Stro	ke index	38.6±12.21	33.3±8.42	35.2±6.63	46.9±18.02	43.2±13.87	35.8±6.84	
Cardiac	M-mode	5.76±1.76	5.0±1.27	5.09±0.74	7.92±2.42	7.18±2.16	6.12±1.38	
Output	Lvot method	5.68±1.59	5.02 ±1.27	5.05±0.87	7.97±2.35	7.11±2.28	6.30±1.64	
	liac index : method	3.60 ±1.10	3.21±0.75	3.24 ±0.57	5.0 ±1.43	4.55±1.37	3.7±0.93	
Distensibility index of lvc		0.63±0.13	0.73±0.14	0.75±0.11	0.46±0.15	0.43±0.13	0.63±0.17	
He	Heart rate		95 ±10	93±8	112± 16	101±24	104± 6	

The above table shows serial echocardiographic measurements of study subjects in terms of survival and non survival. LVIDD was overall in the normal range in survivor and they were generally lower than the non survivors. While in non survivors, LVIDD initially in the normal range which gradually started increasing. similarly LVIDS was overall in the normal range in the patients of sepsis who survived. While in non survivors, LVIDS initially in the normal range which gradually started dilating. Ejection Fraction was around 50% in the survivors which was maintained throughout their monitoring. However in non survivors, ejection fraction

was initially higher than survivors which was gradually decreased. Stroke volume, stroke index, Cardiac output and cardiac index were initially higher in survivor group at the time of admission which was gradually decreased and again started increasing maintaining normal range. While in non survivor, Stroke volume, stroke index, Cardiac output and cardiac index were initially higher on admission and more than survivor which were gradually decreased but remains in higher range throughout their monitoring. Heart rate was in the higher range in the non survivors than survivors throughout their monitoring.

Table 3: Comparative echocardiographic data between sepsis, severe sepsis and septic shock

	SEPSIS	<u> </u>		E SEPSIS		SEPTIC SHOCK	
	S(n=26)	NS	S(n=18)	NS(n=14)	S(n=2)	NS(n=10)	
	45.2±5.7		49.3±7.7	45.3±7.5	38±8.5	49.4±3.7	
LVIDD	43.8±4.0		45.4±3.3	47.7 ±5.6*	42±5.6	47.4±2.4●	
	44.9 ±3.6		44.3±3.1	49.8±5.5®	47.5±0.7	50±3.46®	
	31.3±6.1		36.6±9.0	30.4±6.2	31±7.1	33.3±4.2	
LVIDS	32.4±5.0		31.8±4.9	34.7±4.3	29.5±7.8	33.7±3.6	
	33.7±3.4		31.7±4.7	38.8±3.3	33.5±0.7	37.3±9.8	
	52± 15		48±14	56±13	37±5	57±11	
EF	49±13		52±14	46±7	54±16	50±15	
	47±8		51±11	35±8	50±1	51±25	
	58.3±18.5		66.3±19.1	69.9±30.6	28±15.5	80.3±23.9	
SV	49.6±12.5		57.4±14.4	70±24.7	47.5±10.6	66.6±19.8	
	53.8±10.9		55±9.36	52.3±14.3	69.5±2.1	67±12.5	
	37.5± 10.67		42.6±12.28	44.4±19.96	17.5±7.21	50.5±15.2	
SI	32.2±7.13		35.3±10.3	44.7±16.6	30.5±1.97	41.6±10.8	
	34.6±6.10		35.1±7.29	34.4±7.61	41.8±3.25	37.9±6.5	
	5.86±2.01		5.84±1.35	7.89±2.66	3.8±1.13	7.98±2.18	
CO-M-MODE	4.85±1.25		5.18±1.38	7.53±2.47	5.4±0.28	6.8±1.81	
	4.95±0.80		5.15±0	5.55±1.51	6.25±0.07	6.9±0.87	
	5.72± 1.76		5.86±1.28	7.80±2.45	3.65±0.21	8.22±2.30	
CO LVOT	4.80±1.37		5.32±1.12	7.25±2.51	5.15±0.92	7.02±2.12	
	4.98±0.91		4.99±0.66	5.50±1.29	6.45±1.48	7.27±1.72	
	3.62±1.18		3.72±0.85	4.94±1.52	2.4±0.14	5.16±1.35	
CILVOT METHOD	3.11±0.75		3.38±0.78	4.59±1.59	3.25±0.07	4.39±1.14	
	3.23±0.59		3.17±0.50	3.23±0.66	4.05±0.35	4.40±0.87	
DI	0.67±0.11		0.59±0.13	0.45±0.16	0.4±0.14	0.46±0.14	
	0.74±0.13		0.71±0.15	0.39±0.07	0.61±0.02	0.48±0.18	
	0.77±0.11		0.72±0.11	0.73±0.07	0.75±0.07	0.50±0.17	
	100±10		95±14	118±18	109±16	103±8	
HR	96±8		92±11	98±33	113±10	104±5	
	92±4		93±10	99±2	99±16	110±2	

^{*}n=10, *n=4, •n=9, *n=3.

The above table shows comparative data between sepsis, severe sepsis and septic shock patients and their survival. In the patient of sepsis, there were no significant difference during monitoring, only cardiac output and cardiac index were initially increased which came to normal. In patient of severe sepsis who survived, there were initial increase in cardiac output and LVIDD and cardiac index which gradually returns to normal range. While in non survival of severe sepsis, there were initial increase in ejection fraction, cardiac output and cardiac index which gradually decreased but were significantly more than survival. LVIDD and LVIDS were initially in normal range which gradually increased their dimensions. In patient of septic shock who survived, there were initial decrease in ejection fraction, cardiac output and LVIDD and cardiac index which gradually returns to normal range. While in non survival of septic shock, there were initial increase in ejection fraction, cardiac output and cardiac index which were remain in the higher range throughout monitoring and were significantly more than survival.

 Table 4: Echocaediographic indices in patient who died

Type of Patients	<2 Days	3-5 days	>5 days
High CO and high EF(Distributive Shock)	3	5	1
High CO and low EF (Distributive Shock)	1	3	4
Low CO and low EF (Cardiogenic Shock)	1	4	2
Total	5	12	7

There were 5 patients who died within 48 hours. Of the 5 patients, 3 patients had high cardiac output and high ejection fraction, 1 patient had High Cardiac Output and Low Ejection Fraction (Distributive Shock). 1 had Low Cardiac Output and Low Ejection Fraction (Cardiogenic Shock). There were 12 patients who died within 3-4 days. Of the 12 patients, 5 patients had high cardiac output and high ejection fraction. 3 patients had High Cardiac Output and Low Ejection Fraction (Distributive Shock). 4 had Low Cardiac Output and Low Ejection Fraction (Cardiogenic Shock). There were 7 patients who died after 5 days. Of the 7 patients, 1 patients had high cardiac output and high ejection fraction. 4 patients had High

Cardiac Output and Low Ejection Fraction (Distributive Shock) 2 had Low Cardiac Output and Low Ejection Fraction (Cardiogenic Shock). Thus 17 patients died with Distributive Shock (71%) and 7 patients (29%) died with Cardiogenic Shock had myocardial depression.

Table 5: Multiple logistic regression showing association of various clinical parameters with outcome of sepsis

С	linical	Odds	95% C. I.	'z'	ʻp'
Par	ameters	Ratio	95% C. I.	value	value
	Age	0.93	0.85 to 1.03	-1.43	0.152
	RR	0.39	0.16 to 0.68	-2.97	0.003
	TLC	1.00	1.003to1.004	2.27	0.023
PLA	ATELETS	1.00	1.0 to 1.0001	2.49	0.013
G	RADES	0.04	0.003 to 0.46	-2.57	0.01
HEA	RT RATE	0.47	0.25 to 0.91	-2.25	0.024
	EF	0.88	0.73 to 1.06	-1.29	0.195
	SI	0.36	0.15 to 0.86	-2.29	0.022
	CO	0.08	0.004 to 1.72	-1.60	0.109
	CI	5812	3.62 to 9.31	2.22	0.026

The table shows association of various clinical parameters on admission with outcome of sepsis in multiple logistic regression analysis. In this model, age and grades of sepsis are taken as categorical variables. The analysis shows that after multiple logistic regression, RR (p<0.003, OR=0.39, 95% CI =0.16 to 0.68), TLC(p<0.023, OR=1.00, 95% CI =1.003 to 1.004), PLATELETS(p<0.013, OR=1.00, 95% CI =1.0 to 1.0001), GRADES (p<0.001, OR=0.04, 95% CI =0.03 to 0.46),HEART RATE(p<0.024, OR=0.47, 95% CI =0.25 to 0.91),SI(p<0.022, OR=0.36, 95% CI =0.15 to 0.86),and CI(p<0.026, OR=5812, 95% CI =3.62 to 9.31)are found to be independently associated with outcome of sepsis.

DISCUSSION

Age And Sex: The mean age of the study subjects was 43.2 ± 15.3 years with majority of the study subjects being in the age group of 41-50 years. There was a predominant involvement of the male sex (71%), with females comprising 29% of the study group.

Echocardiographic indices in study subjects: Parker et al^{27} In 1984, performed serial RNCA and simultaneous thermodilution cardiac output studies on 20 patients with septic shock. The 13 survivors and 7 nonsurvivors had the typical hyperdynamic (high cardiac output, low SVR) profile of septic shock. In our study, Comparative echocardiographic study between sepsis, severe sepsis and septic shock patients and their survival shows that in the patient of sepsis, there were no significant difference during monitoring, only cardiac output and cardiac index were initially increased which came to normal. In patient of severe sepsis who survived, there were initial increase in cardiac output and LVIDD and cardiac index which gradually returns to normal range. While in non survival

of severe sepsis, there were initial increase in ejection fraction, cardiac output and cardiac index which gradually decreased but remained higher than the survivors. LVIDD and LVIDS were initially in normal range which gradually increased their dimensions. In patient of septic shock who survived, there were initial decrease in ejection fraction, cardiac output and LVIDD and cardiac index which gradually returns to normal range. While in non survival of septic shock, there were initial increase in ejection fraction, cardiac output and cardiac index which were remain in the higher range throughout monitoring and were significantly more than survival. Thus in our study 17 patients died with distributive shock (71%) and 7 patients (29%) died with cardiogenic shock had myocardial depression.

Effect of various clinical and echocardiographic indices on outcome of sepsis: Cornelis PC de Jager et al^{21} in 2010, studied 92 patients of sepsis to evaluate the ability of conventional infection markers, lymphocyte count and NLCR to predict bacteraemia in adult patients admitted to the Emergency Department with suspected community acquired bacteraemia. They found significant differences between patients with positive and negative blood cultures were detected with respect to the CRP level (mean \pm standard deviation 176 \pm 138 mg/l vs. 116 \pm 103 mg/l; P = 0.042), lymphocyte count (0.8 \pm 0.5 \times $109/l \text{ vs. } 1.2 \pm 0.7 \times 109/l; P < 0.0001) \text{ and NLCR } (20.9)$ \pm 13.3 vs. 13.2 \pm 14.1; P < 0.0001) but not regarding WBC count and neutrophil count. In multivariate analysis adjusted for age and severity of infection, only following admission parameters are associated with negative outcome of sepsis (mortality) viz high respiratory rate, high TLC, low platelets, grades of sepsis, high heart rate. stroke index and cardiac index are found to be independently associated with sepsis.

CONCLUSIONS

Of the 70 patients included in the study, 46 (66%) survived and 24 (34%) succumbed to the illness. The mortality was more in the septic shock patients (83%) as compared to those with severe sepsis (43.7%). There was no mortality in patients with sepsis without any organ involvement. Clinical parameters respiratory rate, platelets, TLC adjusted for age and severity of infection are found to be independently associated with outcome of sepsis in univariate as well as multivariate analysis. Ejection Fraction and Cardiac Output was higher in Non survivors than Survivors at time of admission. However in non survivors, there were progressive decline in ejection fraction over a period of stay but still their cardiac output was in normal range (Distributive Shock). Few had low cardiac output and low ejection fraction when they succumbed (Cardiogenic Shock).¹⁷ (71%)

patients had distributive shock before their death and 7 (29%) had cardiogenic shock and myocardial depression before their death.

REFERENCES

- Annane D, Bellissant E, Cavaillon JM (2005) Septic shock. Lancet 365: 63–78
- Martin GS, Mannino DM, Eaton S, Moss M (2003) ;The epidemiology of sepsis in the United States from 1979 through 2000. N Engl J Med 348: 1546–1554.
- Angus DC, Linde-Zwirble WT, Lidicker J, Clermont G, Carcillo J, Pinsky MR (2001) Epidemiology of severe sepsis in the United States: analysis of incidence, outcome, and associated costs of care. Crit Care Med 29: 1303–1310.
- Rivers E, Nguyen B, Havstad S, Ressler J, Muzzin A, Knoblich B, Peterson E, Tomlanovich M; Early Goal-Directed Therapy Collaborative, Group (2001) Early goal-directed therapy in the treatment of severe sepsis and septic shock. N Engl J Med 345: 1368–1377
- Kern JW, Shoemaker WC (2002) Meta-analysis of hemodynamic optimization in high-risk patients. see comment. Crit Care Med 30: 1686–1692
- Gibot S. Kolopp-Sarda MN. Bene MC. Cravoisy A. Levy B. Faure GC. Bollaert PE (2004) Plasma level of a triggering receptor expressed on myeloid cells-1: its diagnostic accuracy in patients with suspected sepsis. Annals of Internal Medicine. 141:9–15
- Meisner M (2005) Biomarkers of sepsis: clinically useful? Current Opinion Crit Care 11: 473–480
- Farand P, Hamel M, Lauzier F, Plante GE, Lesur O (2006) Review article: organ perfusion/permeabilityrelated effects of norepinephrine and vasopressin in sepsis. Can J Anaesth 53: 934–946
- 9. Pfeiffer; Untersuchungeniiber das choleagift. Ztschr.f.hyg.1891-92, 11; 393-412
- Centanni; Untersuchungeniiber das in fektionsfieber das fieber gift BakterienDtsch med Wochenschr 1894,20;148
- 11. Bone RC. Balk RA. Cerra FB. Dellinger RP. Fein AM. Knaus WA. Schein RM. Sibbald WJ (1992) American College of Chest Physicians/Society of Critical Care Medicine Consensus Conference: definitions for sepsis and organ failure and guidelines for the use of innovative therapies in sepsis. Crit. Care Med 20: 864–874
- 12. Levy MM, Fink MP, Marshall JC, Abraham E, Angus D, Cook D, Cohen J, Opal SM, Vincent JL, Ramsay G,

- International Sepsis Definitions C (2003) 2001 SCCM/ESICM/ACCP/ ATS/SIS International Sepsis Definitions Conference. Intensive Care Med 29: 530–538
- 13. Calandra T, Cohen J, International Sepsis Forum Definition of Infection in the ICU Consensus, Conference (2005) The international sepsis forum consensus conference on definitions of infection in the intensive care unit. Crit. Care Med 33: 1538–1548
- Antonelli M, Levy M, Andrews PJ, Chastre J, Hudson LD, Manthous C, Meduri GU, Moreno RP, Putensen C, Stewart T, Torres A (2007) Hemodynamic monitoring in shock and implications for management: International Consensus Conference, Paris, France, 27–28 April 2006. Intensive Care Med 33:575–590.
- Ailko W. J. Bossink, MD; A. B. Johan Groeneveld, MD, PhD; C. Erik Hack, MD, PhD; and Lambertus G. Thijs, MD, PhD The Clinical Host Response to Microbial Infection in Medical Patients With Fever; Chest 1999;116;380-390.
- Parker SM, Shelhamer JH, Natanson C, Alling DW, Parrillo JE: Serial cardiovascular variables in survivors and non-survivors of human septic shock: heart rate as an early predictor of prognosis. Crit Care Med 1987, 15:923-929.
- Parker MM, Shelhamer JH, Bacharach SL, Green MV, Natanson C, Frederick TM, Damske BA, Parrillo JE: Profound but reversible myocardial depression in patients with septic shock. Ann Intern Med 1984, 100:483-490.
- Amrs.omar.,masudur. Rehman;left atrial function for outcome prediction in severe sepsis and septic shock-an echocardiographic study Indian journal of crit..care Med april-june 2009 vol13
- 19. Silva E, Pedro M D A, Sogayar A C B, Silva T M D O, Janiszewski M Rodrigues R G *et al*, Brazilian sepsis epidemiological study". The critical care forum2004; 8(4): 251-60
- Baumgartner J, Vaney C, Perret C: An extreme form of hyperdynamic syndrome in septic shock. Intensive Care Med 1984, 10:245-249.
- Cornelis PC de Jager1, Paul TL van Wijk, Rejiv B Mathoera, Jacqueline de Jongh-Leuvenink, Tom van der Poll, Peter C Wever Lymphocytopenia and neutrophillymphocytecount ratio predict bacteremia better thanconventional infection markers in an emergencycare unit Critical Care 2010, 14:R192

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