ORIGINAL RESEARCH

A study of etiology, clinical features, and outcome in patients with Multi Organ Dysfunction Syndrome

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ABSTRACT

Background: Multiple organ Dysfunction Syndrome (MODS) is defined as the acute and potentially reversible dysfunction of two or more organs triggered by multiple clinical or non-clinical factors. There is a need to identify the common infections that leads to mortality in intensive care unit. Studies in India have focussed on patients with sepsis due to established causes. Aim and Objective: To know clinical presentation. Etiology and outcomes in patients with multiple organ dysfunction syndrome. Materials and Method : This was hospital based prospective observational study conducted in department of general medicine, Guntur Medical College, GGH, Guntur, for period of two years, in which 100 individuals admitted in ICU with multi organ dysfunction were included in the study. Results: 86% of the patients were form the age group of 31 - 70years and among that also maximum patients were from 61-70 years of age, 48% of the patients were hypertensive and 32% were diabetic. 87% of the patients were admitted with fever and also with other clinical presentation. , 61% had LRTI, 9% had Bacteremia, 12% had Dengue, 5% had pancreatitis, 2% had Congestive heart failure, 6% had renal failure, 5% had hepatic failure. Among all the study population mortality due to MODS was observed among 26% of the patients. SOFA Score, Duration of ICU, Duration Hospital stay and ventilation use were more among non-survivals compared to survivals. Conclusion: Increase in the severity of organ dysfunction which was assessed by SOFA score is very much associated with higher mortality. Etiology of multiple organ dysfunction syndrome can be influenced by regional and seasonal tends. Rickettsial fever, leptospirosis and dengue are common causes of undifferentiated fever in patients with MODS.

Keywords: Multiple organ Dysfunction Syndrome, sequential organ failure assessment, ICU, LRTI

INTRODUCTION

Multiple organ Dysfunction Syndrome (MODS) is defined as the acute and potentially reversible dysfunction of two or more organs triggered by multiple clinical or non-clinical factors. The concept of MODS was first proposed in 1992, which was previously known as multiple organ failure (MOF).^[1] Given that MOF could only be described statically, without showing a continuous process of multiple organ dysfunction, the concept of MODS came into being and gradually replaced MOF.^[2] The organ or system most easily affected by MODS successively include lung, cardiovascular system, liver, kidney, blood system, gastrointestinal tract, and central nervous system. The mortality in MODS patients

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increases with the number of organs involved. When only two organs become dysfunctional, the mortality is about 30%; when 3 or 4 organs are affected, the mortality will rise to 50-70%.^[3, 4]

Two primary causes of MODS are infectious and non-infectious factors, especially the former. Specifically, common causes, in addition to the most common cause of sepsis, include trauma, burn, surgery, shock, and so on.^[3,5] Although an underlying pathophysiology for MODS remains elusive, global perfusion deficits,^[6] widespread endothelial damage,^[7] mitochondrial dysfunction/hibernation and associated energy deficit,^[8] intestinal bacterial product translocation,^[9] and apoptosis^[10] have been implicated. These pathological mechanisms may aggravate the dysfunction of various organs.^[11–14]

In India infections causing multiple organ dysfunction leads to burden of sepsis in ICU. Sepsis can be reversible but as it progresses to septic shock mortality rate increase substantially. Majority of the patients present with clinical features such as fever with myalgia, fever with arthralgia, fever with icterus, fever with rash, or acute encephalitis. Due to their varied presentation, multi system involvement, and lack of sensitivity tests to identify these infections add to diagnostic dilemma.

There is a need to identify the common infections that leads to mortality in intensive care unit. Studies in India have focussed on patients with sepsis due to established causes eg;malaria, leptospira, rickettsial infections. very few studies were done to study the clinical course of the disease in patients with acute undifferentiated fever.

Thus we have undertaken this study to know clinical presentation, Etiology and outcomes in patients with multiple organ dysfunction syndrome

MATERIAL AND METHODS

This was hospital based prospective observational study conducted in department of general medicine, Guntur Medical College, GGH, Guntur, for period of two years, in which 100 individuals admitted in ICU with multi organ dysfunction were included in the study after following inclusion and exclusion criteria and getting informed consent and approved by institutional ethical committee of our Institute.

Inclusion Criteria

• Willing patients with MODS defined as the presence of altered organ function in two or more organ systems.

Exclusion Criteria

- Patients not willing for study
- Patients below 18 years of age
- Do not resuscitate state
- Patients with immunosuppressive state including malignancy and HIV infection, organ transplantation
- Patients with chronic underlying diseases such as CKD,CLD
- Congenital & acquired heart diseases
- SARS CoV-2 Infection

Method

Patients admitted in ICU were included in the study, detailed history was obtained from patient/relative regarding onset duration and progression of symptoms. Complete physical examination was done. Baseline investigations was done for every patient. Additional investigations including those for ruling out cause of fever, imaging studies (xray, ultrasound) was done based on indication for individual patient. Daily investigations were recorded and the progress of patient was assessed by sequential organ failure assessment score (SOFA) till discharge or death or a maximum period of 14 days. The outcome in terms of morbidity and mortality was documented, etiology was identified based on serology and culture.

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The patients were divided into subgroups for analysis –survivors vs non survivors and were then compared in terms of demographic data, signs, symptoms, clinical course and outcome of the disease.

Statistical Analysis

Collected data were entered in the Microsoft excel 2016 for further statistical analysis. Qualitative data were presented in frequency and proportion while quantitative data were expressed as mean and standard deviation. Mean difference of quantitative data between the groups were assessed by using t-test. P-value<0.05 considered as statistically significant. Statistical analysis was done by using statistical software SPSS version 25.

RESULTS

Overall in the study we have included 100 patients with MODS in ICU, there demographic distribution shown in bellow table 1.

Table 1: Distribution of demographic profile among study population				
Parameters	Frequency	Percentage		
Age (Years)				
21 - 30	8	8		
31-40	14	14		
41 - 50	22	22		
51-60	24	24		
61 – 70	26	26		
71-80	6	6		
Mean ± SD	51.85 ± 13.12 Years			
Gender				
Male	52	52		
Female	48	48		
Comorbid Condition				
Diabetes Mellitus	32	32		
Hypertension	48	48		

It was observed that majority (86%) of the patients were form the age group of 31 - 70 years and among that also maximum patients were from 61-70 years of age, and mean age of the patients was more than 50 years. Also among population male predominancy was observed compared to female. And overall 48% of the patients were hypertensive and 32% were diabetic.

 Table 2: Clinical presentation and GCS of study population

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Parameters	Frequency	Percentage		
	Clinical Presentation			
Fever	87	87		
Myalgia	58	58		
Dyspnoea	49	49		
Nausea/Vomiting	49	49		
Cough	56	56		
Arthralgia	51	51		
Headache	48	48		
Abdominal pain	39	39		
Altered sensorium	28	28		

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Loose stools	16	16	
Jaundice	10	10	
Rash	18	18	
Glasgow Coma Scale			
9-12	73	73	
>13	27	27	
Mean ± SD	11.25 ± 1.78		

We have observed 87% of the patients were admitted with fever and also with other clinical presentation like myalgia, cough, arthralgia, dyspnoea, nausea/vomiting, headache, abdominal pain shown as above Table 2.

Table 3 showed distribution of etiology and outcomes of study population, in which we have observed that, 61% had LRTI, 9% had Bacteremia, 12% had Dengue, 5% had pancreatitis, 2% had Congestive heart failure, 6% had renal failure, 5% had hepatic failure. Among all the study population mortality due to MODS was observed among 26% of the patients.

Table 3: Distribution of Etiology and outcome among study population

Parameters	Frequency	Percentage		
Etiology				
LRTI	61	61		
Bacteraemia	9	9		
Dengue	12	12		
Pancreatitis	5	5		
Congestive heart failure	2	2		
Hepatic failure	5	5		
Renal failure	6	6		
Outcome				
Dead	26	26		
Survived	74	74		

Table 4: Distribution of Etiology and outcome among study population					
Vitals	Survived	Dead	P-value		
Pulse	98.22 ± 15.81	66.53 ± 3.88	0.001*		
SBP	119.43 ± 14.75	71.63 ± 7.80	0.001*		
Respiratory Rate	30.12 ± 3.41	38.03 ± 1.10	0.001*		
Temperature	100.37 ± 0.88	102.3 ± 0.34	0.001*		
SOFA Scale	9.07 ± 2.12	13.83 ± 0.77	0.001*		
ICU Stay	10.37 ± 4.47	21.33 ± 1.41	0.001*		
Duration of fever	9.58 ± 2.25	14.23 ± 0.76	0.001*		
Duration of hospital stay	15.84 ± 4.97	27.53 ± 1.55	0.001*		
Duration of ventilator use	3.69 ± 1.46	6.463 ± 0.51	0.001*		
GCS	12.00 ± 1.44	9.123 ± 0.33	0.001*		

*p-value<0.05, statistically highly significant at 5% level of Significance

From table 4, it was observed that all the vitals were statistically significant between survival and non-survival. Respiratory rate among non-survival was more compared to survivals. SOFA Score, Duration of ICU, Duration Hospital stay and ventilation use were more among non-survivals compared to survivals. Mean Glasgow coma scale was observed moderate among non-survival patients.

DISCUSSION

The incidence of MODS depends on the criterion used for MOF as there is no consensus on a single definition as the gold standard.^[15] The reported incidence of MODS among critically ill trauma patients varies widely from 28% to 88%.^[16] MODS is also considered as the frequent cause of mortality in patients admitted to the ICU and the rate of mortality and length of hospital stay correlated with the number of organs involved and the severity of MODS.^[17] An earlier study reported 15% mortality among high-risk surgical patients admitted to the ICU; of which more than half the patients died primarily due to MOF.^[18] Apart from higher mortality, critically ill patients that developed MODS stayed three times longer in the ICU and necessitated greater mechanical ventilatory support than those without MODS.^[19] To date, various MOF scoring systems have been proposed to assess severity and risk stratification in critically ill patients.^[20] Therefore, it is challenging to compare the incidence of heterogeneous populations using various MODS scoring systems. In addition, there is a lack of consistent data for the course of MODS, the mechanisms of organ dysfunction, and the early prediction of MODS in critically ill patients admitted in different ICUs.

According to a multiyear survey of SICU patients, 54% developed MODS.^[21] It was found that hypoperfusion without shock, sepsis without shock, and shock of any etiology were the most common risk factors for MODS development. The most common cause of death in surgical intensive care units (SICUs) is MODS, and its severity is strongly correlated with mortality and hospitalization length.^[22]

In the present study we have observed that 87% had fever, 58% had myalgia, 49% had dyspnea, 49% had Nausea/Vomiting, 56% had cough, 51% had arthralgia, 48% had headache, 39% had abdominal pain, 28% had Altered sensorium, 16% had Loose stools, 10% had jaundice, 18% had Rash. Study conducted by Bhanukumar Muthaiahet al., observed that fever being universal (100%) among these patients, other common presenting symptoms included vomiting/loose stools (45%) and yellowish discoloration of eyes (30.6%), cough (15%), dyspnea (26.6%)

In the current study we have observed with, in which we have observed that, most common etiology was LRTI(61%) followed by Bacteremia, Dengue, pancreatitis, Congestive heart failure, renal failure, hepatic failure. According to the BhanukumarMuthaiah et al.,[23]most common aetiology for acute febrile illness with MODS was dengue fever in 22 (29.3%) patients followed by leptospirosis in 17(22.7%) patients. one more another study conducted by Desai SR & Lakhani JD et al observed that Cholecystitis (12%), UTI (10%), Meningitis (8%) and Pancreatitis (8%).

In our study we observed mortality among 26% of the patients. According to the BhanukumarMuthaiah et al. patients with central nervous involvement predict highest mortality which was 100% in there study.Desai SR & Lakhani JD et al[24] found, 39 patients of MODS, of which 22 died. Hence the mortality rate of patients who had MODS was 56.4%, they found that overall mortality rate was 48% which was higher compared to other studies. But they had more patient of MODS (78% patient) which might have accounted for higher mortality rate. The mortality rate for sepsis, severe sepsis and MODS was 100%, 10% and 56.4% respectively. they had only one patient of sepsis, which was of burns (85% burns) which might account for false high mortality rate in sepsis group. Otherwise the difference in mortality rate between severe sepsis (10%) group and MODS (56.4%) group was statistically significant (p=0.000). Hence, as the number of organ involved increases, it should give alarming sign to the physician.

In our study we have use sequential organ failure assessment (SOFA) scoring system, to asses the organ failure and we have found that SOFA score was more among non-survival patients compared to survival and this difference in the score was statistically highly

significant (p-value<0.01). also we have found mean ICU stay and duration of hospital stay among non-survival was significantly higher compare with moderate GCS scale compared to survival patients.

CONCLUSION

From overall observation after discussing with various studies we can conclude that, Septic shock at admission is associated with higher mortality. Increase in the severity of organ dysfunction which was assessed by SOFA score is very much associated with higher mortality. Etiology of multiple organ dysfunction syndrome can be influenced by regional and seasonal tends. Rickettsial fever, leptospirosis and dengue are common causes of undifferentiated fever in patients with MODS.

CONFLICT OF INTEREST Nil SOURCE OF FUNDING Self funded

REFERENCES

- 1. Physicians ACoC. 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. (1992) 20:864–74. doi: 10.1097/00003246-199206000-00025
- Beal AL, Cerra FB. Multiple organ failure syndrome in the 1990s. Systemic inflammatory response and organ dysfunction. JAMA. (1994) 271:226–33. doi: 10.1001/jama.271.3.226
- 3. Gourd NM, NikitasN.Multiple organ dysfunction syndrome. J Intensive Care Med. (2020) 35:1564–75. doi: 10.1177/0885066619871452
- 4. Seely AJ, Christou NV. Multiple organ dysfunction syndrome: exploring the paradigm of complex nonlinear systems. Crit Care Med. (2000) 28:2193–200. doi: 10.1097/00003246-200007000-00003
- 5. Alam A, Hana Z, Jin Z, Suen KC, Ma D. Surgery, neuroinflammation and cognitive impairment. EBioMedicine. (2018) 37:547–56. doi: 10.1016/j.ebiom.2018.10.021
- 6. Wong BT, Chan MJ, Glassford NJ, Mårtensson J, Bion V, Chai SY, et al. Mean arterial pressure and mean perfusion pressure deficit in septic acute kidney injury. J Crit Care. (2015) 30:975–81. doi: 10.1016/j.jcrc.2015.05.003
- Barichello T, Generoso JS, Singer M, Dal-Pizzol F. Biomarkers for sepsis: more than just fever and leukocytosis-a narrative review. Crit Care. (2022) 26:14. doi: 10.1186/s13054-021-03862-5
- Zou M, Su X, Wang L, Yi X, Qiu Y, Yin X, et al. The molecular mechanism of multiple organ dysfunction and targeted intervention of COVID-19 based on timeorder transcriptomic analysis. Front Immunol. (2021) 12:729776. doi: 10.3389/fimmu.2021.729776
- 9. Assimakopoulos SF, Triantos C, Thomopoulos K, Fligou F, Maroulis I, Marangos M, et al. Gut-origin sepsis in the critically ill patient: pathophysiology and treatment. Infection. (2018) 46:751–60. doi: 10.1007/s15010-018-1178-5
- Linkermann A, Stockwell BR, Krautwald S, Anders HJ. Regulated cell death and inflammation: an auto-amplification loop causes organ failure. Nat Rev Immunol. (2014) 14:759–67. doi: 10.1038/nri3743
- 11. Bone RC. Immunologic dissonance: a continuing evolution in our understanding of the systemic inflammatory response syndrome (SIRS) and the multiple organ

dysfunction syndrome (MODS). Ann Inter Med. (1996) 125:680-7. doi: 10.7326/0003-4819-125-8-199610150-00009

- Marshall JC. Inflammation, coagulopathy, and the pathogenesis of multiple organ dysfunction syndrome. Crit Care Med. (2001) 29(7. Suppl):S99– 106. doi: 10.1097/00003246-200107001-00032
- 13. Aird WC. The role of the endothelium in severe sepsis and multiple organ dysfunction syndrome. Blood. (2003) 101:3765–77. doi: 10.1182/blood-2002-06-1887
- 14. Gando S. Microvascular thrombosis and multiple organ dysfunction syndrome. Crit Care Med. (2010) 38(2 Suppl):S35-42. doi: 10.1097/CCM.0b013e3181c9e31d
- Sauaia A, Moore EE, Johnson JL, Ciesla DJ, Biffl WL, Banerjee A. Validation of postinjury multiple organ failure scores. Shock. 2009 May;31(5):438–47. doi: 10.1097/SHK.0b013e31818ba4c6
- 16. Dharap SB, Ekhande SV. An observational study of incidence, risk factors & outcome of systemic inflammatory response & organ dysfunction following major trauma. Indian J Med Res. 2017 Sep;146 (3):346–53. doi: 10.4103/ijmr.IJMR_1538_15
- 17. Barie PS, Hydo LJ. Influence of multiple organ dysfunction syndrome on duration of critical illness and hospitalization. Arch Surg1996;131:1318–24.
- Lobo SM, Rezende E, Knibel MF, Silva NB, Páramo JA, Nácul FE, et al. Early determinants of death due to multiple organ failure after noncardiac surgery in highrisk patients. AnesthAnalg. 2011 Apr;112(4):877– 83. doi: 10.1213/ANE.0b013e3181e2bf8e
- 19. Dewar DC, Tarrant SM, King KL, Balogh ZJ. Changes in the epidemiology and prediction of multiple-organ failure after injury. J Trauma Acute Care Surg. 2013;74 (3):774–9.
- 20. Hutchings L,Watkinson P, Young JD, Willett K. Defining multiple organ failure after major trauma: A comparison of the Denver, Sequential Organ Failure Assessment, and Marshall scoring systems. J Trauma Acute Care Surg. 2017 Mar;82(3):534–41. doi: 10.1097/TA.00000000001328
- 21. Barie PS, Hydo LJ. Epidemiology of multiple organ dysfunction syndrome in critical surgical illness. Surgical infections. 2000 Sep 1;1(3):173-86.
- 22. Barie PS, Hydo LJ. Influence of multiple organ dysfunction syndrome on duration of critical illness and hospitalization. Archives of Surgery. 1996 Dec 1;131(12):1318-24.
- 23. Muthaiah B, Thippeswamy T, Kondareddy S, ChikkegowdaP.Study of Aetiology and Outcome in Acute Febrile Illness Patients with Multiple Organ Dysfunction Syndrome.J Clin of Diagn Res.2016; 10(8):OC16-OC18. <u>https://www.doi.org/10.7860/JCDR/2016/21376/8301</u>
- 24. Desai SR, Lakhani JD: Clinical Profile of Patients with Multi-Organ Dysfunction syndrome in Patients of Sepsis in Rural set up MICU. PJSR2014;7(2):28-32.