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Pediatrics

A Study on Prognostic Indicators of Severe Dengue Fever in Children

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Abstract

Original Research Article

Dengue is the most important emerging tropical viral disease of humans and leading cause of death in SEAR after diarrhea and acute respiratory infections. It is a significant public health concern with its geographical distribution becoming worldwide, involving nearly all the tropical and subtropical regions of the world with significant morbidity and mortality. This study mainly aims at finding out the factors which were associated with severe dengue fever that helps determining the prognosis in children. This case control study was done during November 2017 to October 2018 from the children admitted in Pediatric ward, Vijayanagar institute of medical sciences, Ballari. Among the 201 study subjects, the factors like presence of bleeding manifestations(p<0.001), low pulse pressue<20 mmhg(p<0.001), prolonged capillary refill time(p<0.001), low O2 saturation(p<0.001), Hyponatremia(p=0.008), elevated SGOT/SGPT levels(p<0.001), Prolonged PT/APTT/INR levels(p<0.001), respiratory and abdominal involvement in the form of pleural effusion, pneumonia and ascites(p<0.001) were significantly associated with severe dengue fever with mortality of 15%. Leucopenia, thrombocytopenia and raising hematocrit was not found to have significant association with the severity of dengue fever in this study. These parameters may be used to predict, to forecast disease severity in patients suspected of dengue infection and to manage it early in the course of the illness. Various measures for early diagnosis and prompt intervention with emphasis on the warning signs and good supportive care would help decrease the fatality and morbidity associated with Dengue.

Keywords: Severe dengue fever, thrombocytopenia, hyponatraemia, leucopenia.

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INTRODUCTION

Dengue is an essential mosquito-transmitted viral infection having significant mortality and morbidity. It is a significant public health concern with its geographical distribution becoming worldwide, involving nearly all the tropical and subtropical regions of the world [1]. It is the most common and widespread arboviral infection in the world today caused by Dengue virus. Dengue viruses (DV) belong to the family Flaviviridae, and there are four serotypes of the virus referred to as DV- 1, DV-2, DV-3, and DV-4 [2]. The course of the disease has changed in the recent years from milder form to severe form like dengue hemorrhagic fever & severe dengue fever and with increasing outbreaks frequency [3]. Approximately 1.8 billion (more than 70%) of the population at risk for dengue worldwide live in Member States of the WHO South-East Asia Region (SEAR) and Western Pacific Region, which bear nearly 75% of the current global disease burden due to dengue [4]. During the 1780s, the dengue epidemics were first recognized to occur almost simultaneously in Asia, Africa, and North America, shortly after the identification and naming of the disease

in 1779. Benjamin Rush confirmed the first case report in 1789, and he coined Dengue as "**Breakbone Fever**" because of the symptoms of myalgia and arthralgia [5]. Nearly 10,000 deaths are attributed to dengue infection yearly, which is secondary to rapid urbanization and overcrowding. Temporal trends suggest that the incidence of dengue nearly doubled in every decade since 1990 [6]. Dengue is the primary cause of childhood death in many countries in Southeast Asia [7].

According to WHO 2012 guidelines, the clinical spectrum of disease includes⁸

- Dengue fever
- Dengue with warning signs and
- Severe dengue fever (DF).

Severe dengue is defined as a suspected dengue patient associated with severe bleeding, severe organ dysfunction or severe plasma leakage leading to shock.

In addition to the distinction between severe and non-severe dengue, the WHO recognizes three

Citation: Karthikeyan. A & Sudhakar Hegade. A Study on Prognostic Indicators of Severe Dengue Fever in Children. Sch J App Med Sci, 2021 Jun 9(6): 1013-1028. phases in the clinical course of a dengue infection, i.e. the febrile, critical and recovery phase.

This study is aimed to find out the prognostic indicators in severe dengue fever in children which will help us in turn to identify severe dengue fever at the earliest and to assess the prognosis.

MATERIALS AND METHODS

Data will be collected by face to face interview from the parents of the children admitted in pediatric emergency ward with diagnosis of dengue fever and severe dengue fever in Department of pediatrics, Vijayanagar institute of medical sciences, Ballari from November 2017 to October 2018. Clinical parameters will be examined and documented.

Method of Collection of Data

This is a case control study design in which 201 cases of dengue fever admitted in the pediatric emergency ward, were enrolled. After taking written informed consent, data were collected in a predesigned semi structured questionnaire regarding Socio-Demographic profile, medical history, clinical and hematological profile and outcome. Investigations such as Complete blood count, Blood test(Serology) for Dengue fever, urine routine, serum electrolytes, random blood sugar, WIDAL/ Blood culture, Liver enzymes, Chest x ray, Ultrasound abdomen, PT, aPTT, INR were done based on the clinical scenario and if necessary, blood urea, serum creatinine, CSF analysis, 2D echo, CT scan brain/MRI was also done. Non randomized purposive sampling technique will be adopted to select both cases and controls.

Sample Size Calculation

Since this study is a time bound study, where a series of dengue cases (as per inclusion criteria) were enrolled during a study period of one year i.e., from November 2017 to October 2018. During this study period, we were able to enroll a total of 201 cases, out of which 142 were only dengue fever, 32 were dengue fever with warning signs and 27 were severe dengue fever.

Inclusion Criteria

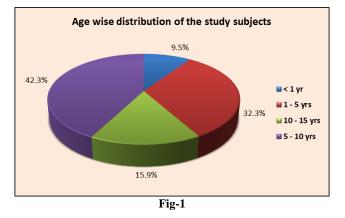
- I. For cases: Any child, between age 1 month to 12 years, having tested positive for dengue NS1 antigen in serum or tested positive for antibodies (IgM) in serum against dengue fever virus along with complications such as shock or fluid accumulation with respiratory distress or severe bleeding or severe organ involvement such as liver, CNS, heart and other organs.
- II. **For controls:** Any child, between age 1 month to 12 years, having tested positive for dengue NS1 antigen in serum or tested positive for antibodies (IgM) in serum against dengue fever virus without any complications.

Exclusion Criteria

- 1. Children having other co-infections like malaria, typhoid or infective hepatitis, interfering with interpretation of the laboratory data.
- 2. Immunocompromised subjects.
- 3. Children less than 1 month.
- 4. Patients discharged against medical advice.

RESULTS

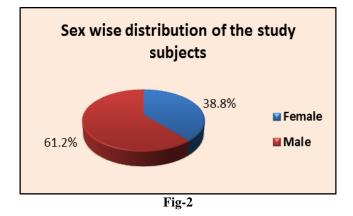
Table-1					
Age wise dis	tribution of th	e study subjects			
Age group	Frequency	Percent			
< 1 yr	19	9.5			
1 - 5 yrs	65	32.3			
5 - 10 yrs	85	42.3			
10 - 15 yrs	32	15.9			
Total	201	100			
$Mean \pm SD$	6.59 ± 3.96	Mean \pm SD			



Among the 201 cases included in the study, 19(9.5%) were less than 1 yr, 65(32.3%) cases were 1-5

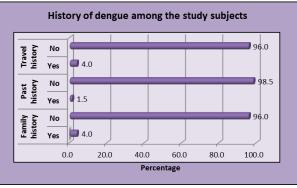
yrs, 85(42.3%) were between 5-10 yrs and 32(15.9%) were between 10-15 yrs of age.

	Table-2		
Sex wise distribution of the study subjects			
Sex	Frequency	Percent	
Female	78	38.8	
Male	123	61.2	
Total	201	100	



78(38.8%) cases were females and 123(61.2%) cases were males.

	Table-3					
History of dengue among the study subjects (n=201)						
History of dengue Frequency Percentage						
Family history						
Yes	8	4.0				
No	193	96.0				
Past history						
Yes	3	1.5				
No	198	98.5				
Travel history						
Yes	8	4.0				
No	193	96.0				



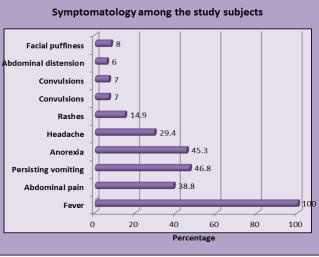


8(4%) of cases had positive family h/o dengue fever, 3(1.5%) cases had positive past h/o dengue fever

and 8(4%) cases had positive h/o travel to dengue endemic areas.

Table-4						
Symptomatology among the study subjects (n=201)						
Symptoms	Frequency	Percentage				
Fever	201	100				
Abdominal pain	78	38.8				
Persisting vomiting	94	46.8				
Anorexia	91	45.3				
Headache	59	29.4				
Rashes	30	14.9				
Convulsions	14	7.0				
Abdominal distension	12	6.0				
Facial puffiness	16	8.0				







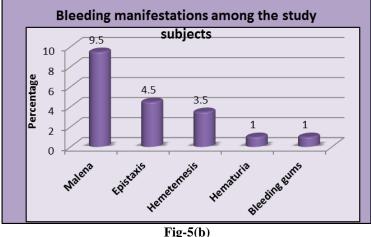
201(100%) cases had fever, 94(46.8%) had persistent vomiting, 91(45.3%) had anorexia, 78(38.8%) had abdominal pain, 59(29.4%) had headache, 30(14.9%) had rashes, 16 (8%) had facial puffiness, 14(7%) had convulsions and 12(6%) had abdominal distension.

Table-5(a)						
Distribution of study subjects based on bleeding manifestation						
Bleeding MF Frequency Percent						
Yes	31	15.4				
No	170	84.6				
Total	201	100				

Table-5(b)						
Bleeding manifestations among the study subjects (n=201)						
Bleeding MF	Frequency	Percentage				
Malena	19	9.5				
Epistaxis	9	4.5				
Hemetemesis	7	3.5				
Hematuria	2	1				
Bleeding gums	2	1				

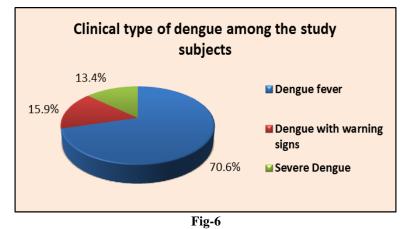
Bleeding manifestations were seen in 31(15.4%) of cases with 19(9.5%) had melena, 9(4.5%) had epistaxis, 7(3.5%) had hemetemesis and 2(1%) had

hematuria and bleeding gums. 170(84.6%) cases had no bleeding manifestations.



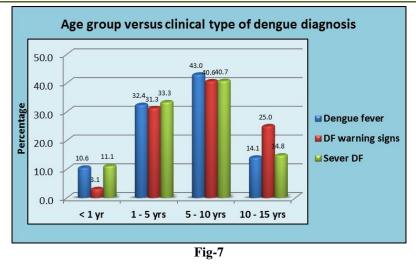
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Table-6						
Clinical type of dengue among the study subjects						
Туре	Frequency	Percent				
Dengue fever	142	70.6				
Dengue with warning signs	32	15.9				
Severe Dengue	27	13.4				
Total	201	100.0				



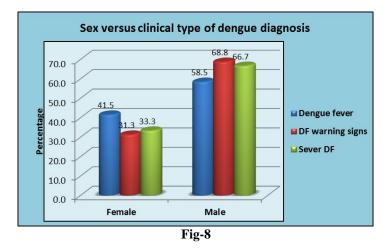
142(70.6%) had dengue fever, 32(15.9%) had dengue fever with warning signs, 27(13.4%) had severe dengue fever.

			Table-'	7			
Age group v	ersus clinical	l type of d	engue diagnos	sis			
Age group	Dengue fev	Dengue fever DF with warning signs Severe DF				P value	
	Frequency	Percent	Frequency	Percent	Frequency	Percent	
< 1 yr	15	10.6	1	3.1	3	11.1	0.71
1 - 5 yrs	46	32.4	10	31.3	9	33.3	
10 - 15 yrs	20	14.1	8	25.0	4	14.8	
5 - 10 yrs	61	43.0	13	40.6	11	40.7	
Total	142	100.0	32	100.0	27	100.0	
Mean \pm SD	6.39 ± 3.96		7.38 ± 4.07		6.67 ± 3.87		0.449



Mean age group of dengue fever were 6.39+3.93 yrs, DF with warning signs were 7.38+4.07 yrs, severe dengue fever were 6.67+3.87 yrs(p=0.449).

	Table-8								
Sex vers	us clinical typ	e of dengu	e diagnosis						
Sex	Sex Dengue fever DF with warning signs Severe DF P v.								
	Frequency	Percent	Frequency	Percent	Frequency	Percent			
Female	59	41.5	10	31.3	9	33.3	0.458		
Male	83	58.5	22	68.8	18	66.7			
Total	142	100.0	32	100.0	27	100.0			

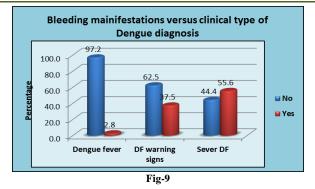


Although in the present study, the incidence of dengue fever, DF warning signs and Severe dengue

fever was more in males, there is no significant association with the severity of dengue fever(p=0.458).

Table-9									
Bleeding main	Bleeding mainifestations versus clinical type of Dengue diagnosis								
Bleeding MF	Dengue fev	Dengue fever DF with warning signs Severe DF							
	Frequency	Percent	Frequency	Percent	Frequency	Percent			
No	138	97.2	20	62.5	12	44.4	< 0.001		
Yes	4	2.8	12	37.5	15	55.6			
Total	142	100.0	32	100.0	27	100.0			

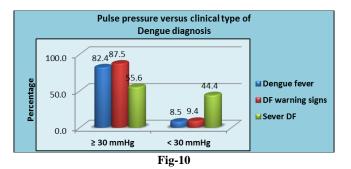
Table 0



The presence of bleeding manifestations like melena, epistaxis, hemetemesis, petechiae, hematuria s more in DF with warning signs(37.5%) and Severe

dengue fever(55.6%). There is significant association between bleeding manifestations and severity of dengue fever (p<0.001).

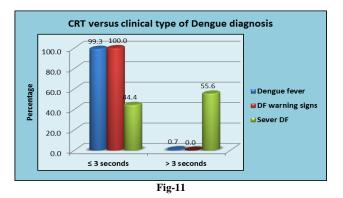
Table-10									
Pulse pressure	Pulse pressure versus clinical type of Dengue diagnosis								
Pulse	Dengue fev	er	DF with war	ning signs	Severe DF		P value		
pressure	Frequency	Percent	Frequency	Percent	Frequency	Percent			
\geq 20 mmHg	117	82.4	28	87.5	15	55.6	< 0.001		
< 20 mmHg	12	8.5	3	9.4	12	44.4			
NR	13	9.2	1	3.1	0	0.0			
Total	142	100.0	32	100.0	27	100.0			



Narrow pulse pressure(<20 mmhg) was found in 8.5% in dengue fever, 9.4% in DF with warning signs, 44.4% in severe dengue fever and it is statistically signific(p<0.001).

Table-11

CRT versus clinical type of Dengue diagnosis										
CRT	Dengue fever		DF with warning signs		Severe DF		P value			
	Frequency	Percent	Frequency	Percent	Frequency	Percent				
\leq 3 seconds	141	99.3	32	100.0	12	44.4	< 0.001			
> 3 seconds	1	0.7	0	0.0	15	55.6				
Total	142	100.0	32	100.0	27	100.0				



Prolonged Capillary refill time of > 3 seconds was found in 55.6% in severe dengue fever which is statistically significant (p<0.001).

	Table-12										
SpO2 levels versus clinical type of Dengue diagnosis											
SpO2	Dengue fev	er	DF with wa	rning signs	Severe DF	P value					
-	Frequency	Percent	Frequency	Percent	Frequency	Percent					
Low	8	5.6	6	18.8	16	59.3	< 0.001				
Normal	134	94.4	26	81.3	11	40.7					
Total	142	100.0	32	100.0	27	100.0					

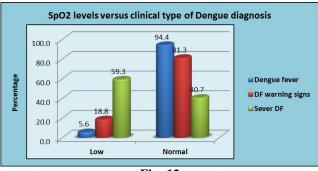
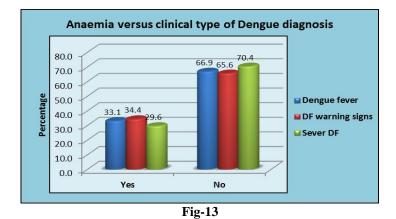


Fig- 12

Low spO_2 were identified in 5.6% in dengue fever, 18.8% in DF with warning signs and 59.3% in

severe dengue fever and this has significant association with severity of dengue fever(p < 0.001).

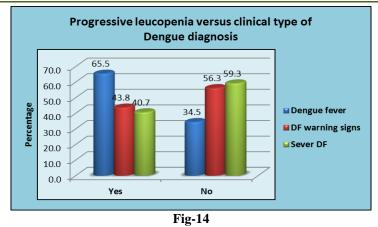
	Table-13											
Anaemia versus clinical type of Dengue diagnosis												
Anaemia	Dengue fev	er	DF with wa	rning signs	Severe DF	P value						
	Frequency	Percent	Frequency	Percent	Frequency	Percent						
Yes	47	33.1	11	34.4	8	29.6	0.921					
No	95	66.9	21	65.6	19	70.4						
Total	142	100.0	32	100.0	27	100.0						



Anaemia was found in 33.1% in dengue fever, 34.4% in DF with warning signs and 29.6% in severe

dengue fever and it is not statistically significant(p=0.921).

	Table-14										
Progressive leucopenia versus clinical type of Dengue diagnosis											
Leucopenia	Dengue fev	er	DF with wa	rning signs	Severe DF	P value					
_	Frequency	Percent	Frequency	Percent	Frequency	Percent					
Yes	93	65.5	14	43.8	11	40.7	0.01				
No	49	34.5	18	56.3	16	59.3					
Total	142	100.0	32	100.0	27	100.0					



Progressive leucopenia was found in 65.5% of dengue fever, 43.8% in DF with warning signs and 40.7% in severe dengue fever and it is not statistically significant in our present study(p=0.01). Leukocytosis

was identified more in severe dengue who presented with/ developed bleeding manifestations which was significantly associated with severity of dengue fever.

Table-15										
Lymphocytopenia versus clinical type of Dengue diagnosis										
Lymphocytopenia	Dengue feve	er	DF with wa	rning signs	Severe DF	P value				
	Frequency	Percent	Frequency	Percent	Frequency	Percent				
Yes	50	35.2	13	40.6	6	22.2	0.306			
No	92	64.8	19	59.4	21	77.8				
Total	142	100.0	32	100.0	27	100.0				

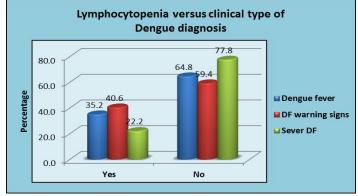
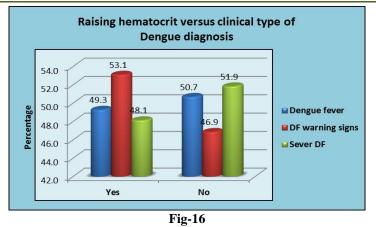


Fig-15

Lymphocytopenia was found in 35.2% in dengue fever, 0.6% in DF with warning signs and

22.2% in severe dengue fever which is not statistically significant.

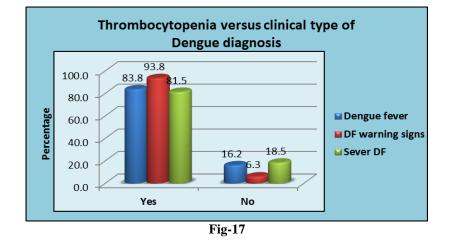
Table-16										
Raising hematocrit versus clinical type of Dengue diagnosis										
Hematocrit	Dengue fev	er	DF with warning signs		Severe DF	P value				
	Frequency	Percent	Frequency	Percent	Frequency	Percent				
Yes	70	49.3	17	53.1	13	48.1	0.912			
No	72	50.7	15	46.9	14	51.9				
Total	142	100.0	32	100.0	27	100.0				



Raising heamtocrit was identified in 49.3% in dengue fever, 53.1% in DF with warning signs and

48.1% in severe dengue fever and it is statistically not significant in our present study(p=0.912).

Table-17										
Thrombocytopenia versus clinical type of Dengue diagnosis										
			DF with	warning						
Thrombocytopenia	Dengue fev	er	signs		Severe DF		P value			
	Frequency	Percent	Frequency	Percent	Frequency	Percent				
Yes	119	83.8	30	93.8	22	81.5	0.308			
No	23	16.2	2	6.3	5	18.5				
Total	142	100.0	32	100.0	27	100.0				



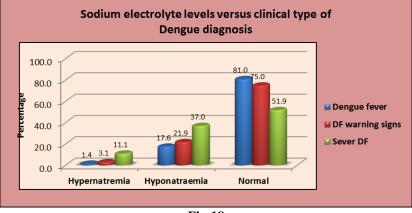
Thrombocytopenia was found in 83.8% in dengue fever, 93.8% in DF with warning signs and

81.5% in severe dengue fever and it is not statistically significant(p=0.308).

	Table-18										
Sodium electrolyte levels versus clinical type of Dengue diagnosis											
Sodium	Dengue feve	e fever DF with warning sign		ning signs	Severe DF	P value					
	Frequency	Percent	Frequency	Percent	Frequency	Percent					
Hypernatremia	2	1.4	1	3.1	3	11.1	0.008				
Hyponatraemia	25	17.6	7	21.9	10	37.0					
Normal	115	81.0	24	75.0	14	51.9					
Total	142	100.0	32	100.0	27	100.0					

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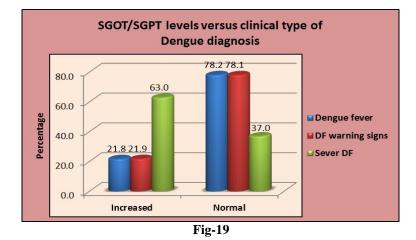




Hyponatraemia was found in 17.6% in dengue fever, 21.9% in DF with warning signs and 37% in

severe dengue fever and this is significantly associated with severity of dengue fever(p=0.008).

Table-19										
SGOT/SGPT levels versus clinical type of Dengue diagnosis										
SGOT/SGPT	Dengue feve	er	DF with warning signs		Severe DF		P value			
	Frequency	Percent	Frequency	Percent	Frequency	Percent				
Increased	31	21.8	7	21.9	17	63.0	< 0.001			
Normal	111	78.2	25	78.1	10	37.0				
Total	142	100.0	32	100.0	27	100.0				

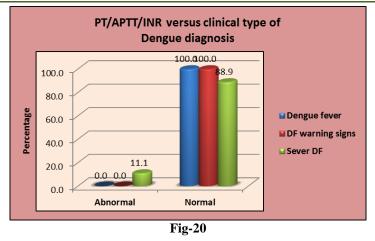


Elevated SGOT/SGPT levels were found in 21.8% in dengue fever, 21.9% in DF with warning signs and 63% in severe dengue fever and this has significant

association with severity of dengue fever(p<0.001). The SGOT levels more than 1500 were having more mortality in our present study.

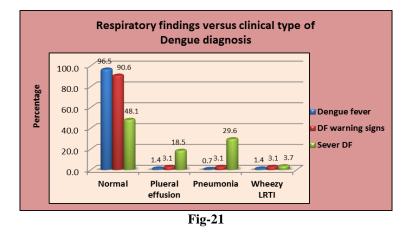
Table-20										
PT/APTT/INR versus clinical type of Dengue diagnosis										
PT/APTT/INR	Dengue feve	er	DF with war	rning signs	Severe DF	P value				
	Frequency	Percent	Frequency	Percent	Frequency	Percent				
Abnormal	0	0.0	0	0.0	3	11.1	< 0.001			
Normal	142	100.0	32	100.0	24	88.9				
Total	142	100.0	32	100.0	27	100.0				

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Prolonged PT/APTT.INR was found 11.1% only in severe dengue fever which is statistically significant (p<0.001).

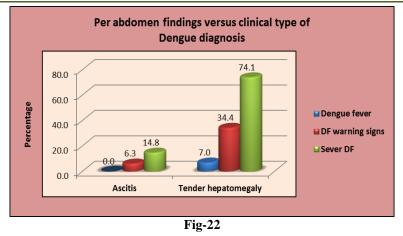
			Table-2	1						
Respiratory findings versus clinical type of Dengue diagnosis										
RS findings	Dengue feve	er	DF with warning signs		Severe DF		P value			
	Frequency	Percent	Frequency	Percent	Frequency	Percent				
Normal	137	96.5	29	90.6	13	48.1	< 0.001			
Plueral effusion	2	1.4	1	3.1	5	18.5				
Pneumonia	1	0.7	1	3.1	8	29.6				
Wheezy LRTI	2	1.4	1	3.1	1	3.7				
Total	142	100.0	32	100.0	27	100.0				



The presence of respiratory system involvement such as respiratory distress, Pneumonia, wheezy LRTI,ARDS and pleural effusion were found in 3.5% in dengue fever, 9.3 in DF with warning signs and 51.8% in severe dengue fever which is statistically significant(p<0.001).

Table-22									
Per abdomen findings versus clinical type of Dengue diagnosis									
Signs	Dengue fever		DF with warning signs		Severe DF		P value		
	Frequency	Percent	Frequency	Percent	Frequency	Percent			
Ascitis	0	0.0	2	6.3	4	14.8	< 0.001		
Tender hepatomegaly	10	7.0	11	34.4	20	74.1	< 0.001		
Total	142	100.0	32	100.0	27	100.0			

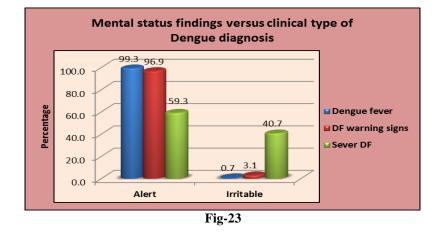
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The presence of ascitis and tender hepatomegaly was found in 7% in dengue fever, 40.7%

in DF with warning signs and 88.9% in severe dengue fever and this is statistically significant(p<0.001).

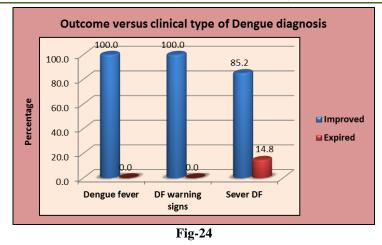
Table-23								
Mental status findings versus clinical type of Dengue diagnosis								
CNS	Dengue fever		DF with warning signs		Severe DF		P value	
	Frequency	Percent	Frequency	Percent	Frequency	Percent		
Alert	141	99.3	31	96.9	16	59.3	< 0.001	
Irritable	1	0.7	1	3.1	11	40.7		
Total	142	100.0	32	100.0	27	100.0		



The presence of altered mental state like irritability was found in 0.7% in dengue fever, 3.1% in

DF with warning sign s and 40.7% in severe dengue fever which is statistically significant(p<0.001).

Table-24									
Outcome versus clinical type of Dengue diagnosis									
Outcome	Dengue fev	er	DF with warning signs		Severe DF		P value		
	Frequency	Percent	Frequency	Percent	Frequency	Percent			
Expired	0	0.0	0	0.0	4	14.8	< 0.001		
Improved	142	100.0	32	100.0	23	85.2			
Total	142	100.0	32	100.0	27	100.0			



The mortality from dengue fever was seen in 14.8% from severe dengue fever.

DISCUSSION

We conducted this study based on the new WHO classification 2012 of dengue fever, to know which factors can predict severity so as to facilitate timely intervention. In our study out of 201 patients with dengue, 142 were classified as dengue fever, 32 as DF with warning signs and 27 children as severe dengue. However Prasad D *et al* found 82.1% as severe dengue and Sahana *et al* found 24.7% [9].

This variance in the subject distribution may be attributable to the timing of admission, the awareness in the draining community, timely intervention.

The mean age of presentation in our study was 6.59 ± 3.96 years with 42.3% of the children in the 5-10 year age group, a finding similar to that by Gomber S *et al* and Rasul *et al* [10, 11].

This could be due to this age group prone for frequent mosquito bites due to their outdoor activities, but our study does not prove any significant association between age and the severity of dengue fever. Studies done by Shah I *et al* and Rachel C *et al* found younger age as one of the predictive markers for Dengue shock syndrome and dengue hemorrhagic fever [12].

The male:female ratio in our study was 1.57:1 similar to study by Rasul *et al*. Though studies by Anders KL *et al* and Phuong CX *et al* found increased risk of shock and death among females, our study did not have a significant statistical association similar to studies by Pongpan S and Ahmed FU *et al* [13].

There was no significant association was found between nutritional status and the severity of dengue fever. Pichainarong stated that severe dengue is rarely seen in children with protein energy malnutrition [14]. Many studies has been done previously to find out the risk factors significantly associated with severity of dengue fever like bleeding manifestations, leucopenia, raising hematocrit, hypotension, narrow pulse pressure, elevated SGOT/SGPT levels, deranged PT/APTT/INR etc. which are similar to our study.

The presence of bleeding manifestations was having a statistically significant association with the severity of dengue fever like many other studies. The most common bleeding manifestations found in our study was melena and epistaxis followed by others.

Narrow pulse pressure(< 20 mmhg) and prolonged Capillary refill time(>3 secs) and altered mental state i.e irritability was found to have significant association with severity of dengue fever as showed in the study done by Rajapakse s *et al* [15].

Low oxygen saturation was found in 59.3 % of severe dengue fever cases in our study, which is showing statistically significant association with the dengue fever severity. Progressive leukopenia, hyponatraemia, elevated SGOT/SGPT levels, prolonged PT/APTT/INR levels, presence of pleural effusion, LRTI, tender hepatomegaly and ascitis was significant association with the severity of dengue fever as shown by many of the studies. But this study did not find the association of anemia, lymphocytopenia, raising hematocrit, thrombocytopenia with severity of dengue fever. This could be because of early identification, diagnosis and intervention with IV fluids resulting in masking of results.

However many studies proposed hemoconcentration as a prognostic factor for the severity of dengue infection [16, 17]. Vasculopathy in dengue causes increased vascular permeability, leading to hemo-concentration and shock. Some studies concluded that white cell count > $5,000/\mu$ L is a prognostic factor for dengue severity [18]. while others found leucopenia. Normoleukocytosis or mild leukocytosis may be found in early dengue infection. When body temperature declines, most patients experienced

leucopenia from bone marrow suppression. Stress in accompanied with shock may somehow cause leukocytosis.

Thrombocytopenia was found to have association with the severity of dengue fever in the studies conducted by Jayashree K *et al*, Mogra G *et al* and Mourao *et al*, but this was not established in our present study [19]. Low platelets are explained by bone marrow suppression and immune response induced platelet destruction by the liver and spleen.

Hyponatraemia was found to have significant association with severity of dengue fever. Studies done by Adisorn Lumpaopong *et al* showed mild hyponatremia is a common electrolyte disturbance and renal involvement is mild in patients with DF and DHF [20].

Studies done by Dayal A *et al* [21] and Sahana *et al* [18] found the association of pleural effusion, ascitis and tender hepatomegaly with the severity of dengue fever, which was again proved in our study. Although moderate liver enlargement is a normal response to dengue infection, it is more associated with severe dengue fever compared to DF.

The mortality among the severe dengue fever in our study was 14.8% which was more than those reported by Sahana *et al* where it was 2%. The overall mortality of dengue fever in our study was 0.02%. Various measures for early diagnosis and prompt intervention with emphasis on the warning signs and good supportive care could help decrease the fatality and morbidity associated with dengue.

CONCLUSION

The severity of dengue infection is significantly associated with some routine clinical parameters like presence of bleeding manifestations, pulse pressue<20 mmhg, prolonged CRT, low O2 saturation, Hyponatraemia, elevated SGOP/SGPT levels, Prolonged PT/APTT/INR levels, respiratory and abdominal involvement. These parameters may be used to predict, to forecast disease severity in patients suspected of dengue infection and to manage it early in the course of the illness. Various measures for early diagnosis and prompt intervention with emphasis on the warning signs and good supportive care would help decrease the fatality and morbidity associated with Dengue.

REFERENCES

 Nathan MB, Drager RD, Guzman M. (2009). Epidemiology, Burden of disease and transmission. In Dengue guidelines for diagnosis, treatment, prevention, and control. WHO: Geneva. p. 1-21

- 2. D. J. Gubler. (1998). Dengue and dengue hemorrhagic fever. Clinl Microbiol Rev, vol. 11, no. 3, pp. 480–496.
- E. Gupta, L. Dar, G. Kapoor, and S. Broor. (2006).
 "The changing epidemiology of dengue in Delhi, India," Virology Journal, vol. 3, article 92.
- Prasittisuk C, Kalra NL, Dash AP. Disease Burden of Dengue Fever and Dengue Haemorrhagic Fever: In Prasittisuk C, Yunnus EB, Moniboth D, Deliana J, Setiabudi D, Dash AP *et al.* editors (2011). In: Comprehensive Guidelines for Prevention and Control of Dengue and Dengue Haemorrhagic Fever. WHO: New Delhi, p. 3-5.
- Dennis Normile. (2013). Surprising New Dengue Virus Throws a Spanner in Disease Control Efforts. Science, 342 (6157): 415. DOI: 10.1126/science.342. 6157. 415).
- 6. Stanaway JD, Shepard DS, Undurraga EA, Halasa YA, Coffeng LE, Brady OJ, *et al.* (2013). The global burden of dengue: an analysis from the global burden of disease study. Lancet Infect Dis, 16:712–23.
- 7. Daumas RP, Passos SR, Oliveira RV, Nogueira RM, Georg I, Marzochi KB, *et al.* (2016). Clinical and laboratory features that discriminate dengue from other febrile illnesses: a diagnostic accuracy study in Rio de Janeiro, Brazil. BMC Infect Dis, 13:77.
- Kliegman RM, Behrman RE, Jenson HB, Stanton BM. (2016). Nelson Textbook of Pediatrics 20e in: Halstead SB, Editors in; Dengue fever and dengue hemorrhagic fever, Elsevier Health Sciences, Philadelphia, 2(3):1629-32.
- Mogra G, Ghildiyal RG, Mohanlal S. (2016). Classification and study of the clinicohematological profile of patients with dengue fever in the pediatric age group. International Journal of Contemporary Pediatrics, 3(4):1405-10.
- Gomber S, Ramachandran VG, Kumar S, Agarwal KN, Gupta P, Dewan DK, *et al.* (2001). Hematological observation as diagnostic marker in dengue hemorrhagic fever- a reappraisal. Indian Pediatr, 38(5):477-81.
- Rasul CH, Ahasan HA, Rasid AK, Khan MR. (2002). Epidemiological factors of dengue hemorrhagic fever in Bangladesh. Indian pediatrics, 39(4):369-72.
- Shah I, Deshpande GC, Tardeja P. (2004). Outbreak of dengue in Mumbai and predictive markers for dengue shock syndrome. Journal of Tropical Pediatrics, 50(5):301-5.
- Pongpan S, Wisitwong A, Tawichasri C, Patumanond J. (2013). Prognostic indicators for dengue infection severity. International Journal of Clinical Pediatrics, 2(1):12-8.
- Pichainarong N, Mongkalangoon N, Kalayanarooj S, Chaveepojnkamjorn W. (2006). Relationship between body size and severity of dengue hemorrhagic fever among children aged 0-14 years.

Southeast Asian J Trop Med Public Health, 37(2):283-8.

- 15. Rajapakse S. (2011). Dengue shock. Journal of Emergencies, Trauma and Shock, 4(1):120.
- Chacko B, Subramanian G. (2007). Clinical, laboratory and radiological parameters in children with dengue fever and predictive factors for dengue shock syndrome. Journal of tropical pediatrics, 54(2):137-40.
- Shah GS, Islam S, Das BK. (2006). Clinical and laboratory profile of dengue infection in children. Kathmandu University medical journal (KUMJ), 4(1):40-3.
- Sahana KS, Sujatha R. (2015). Clinical profile of dengue among children according to revised WHO classification: analysis of a 2012 outbreak from Southern India. The Indian Journal of Pediatrics, 82(2):109-13.

- Mogra G, Ghildiyal RG, Mohanlal S. (2016). Classification and study of the clinicohematological profile of patients with dengue fever in the pediatric age group. International Journal of Contemporary Pediatrics, 3(4):1405-10.
- Lumpaopong A, Kaewplang P, Watanaveeradej V, Thirakhupt P, Chamnanvanakij S, Srisuwan K, Pongwilairat N, Chulamokha Y. (2010). Electrolyte disturbances and abnormal urine analysis in children with dengue infection. Southeast Asian Journal of Tropical Medicine and Public Health, 41(1):72.
- Kittigul L, Pitakarnjanakul P, Sujirarat D, Siripanichgon K. (2007). The differences of clinical manifestations and laboratory findings in children and adults with dengue virus infection. Journal of Clinical Virology, 39(2):76-81.