

Original Research Article

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## Study of Cardiac Manifestations in Dengue Fever

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### ABSTRACT

Dengue fever is an acute febrile infectious disease, caused by any of four serotypes. Severe dengue infections may give rise to many complications such as liver failure, disseminated intravascular coagulation, encephalopathy, myocarditis, acute renal failure and haemolytic uraemic syndrome. ST segment and T wave changes in the electrocardiogram together with low ejection fraction and global hypokinesia on radionuclide ventriculography have been found. Hence, this study was intended to assess the cardiac manifestations in dengue fever patients. To determine the clinical profile of Dengue patients in the study and to study the cardiac manifestations in patients with dengue fever. This study was a hospital based study conducted in Karnataka Institute of Medical Sciences, Hubli. Universal Sample size was taken in the study. I.e. all the cases of Dengue Fever admitted to KIMS Hospital, Hubli for the period of 1 ½ year. All the cases which satisfy the inclusion criteria were included in the study. Informed consent was taken and the clinical and laboratory parameters were recorded in predesigned data sheet. Diagnosis and management were done according to WHO guidelines. Serological tests for detection of IgM dengue antibodies and NS1 antigen was done. All the subjects were subjects for laboratory investigations. Data was entered into Microsoft excel data sheet and was analyzed using SPSS 22 version software. p value (Probability that the result is true) of <0.05 was considered as statistically significant after assuming all the rules of statistical tests. CK-MB and Troponin I at admission were raised in 18% and 72% respectively. On ECG, 56% of the patients had normal rhythm, 15% had sinus Bradycardia, 9% had sinus tachycardia, 10% had T wave changes, 3% had ST Changes, 2% had Sinus Bradycardia with T Wave Changes, 2% had Sinus Tachycardia with T wave changes and 1% had LVH and 2% had Abnormal Rhythm. From this study we can conclude that cardiac manifestations cause significant morbidity and in our study no mortality occurred. Hence cardiac manifestations are important in the management and assessing the prognosis of the patients with dengue.

#### Keywords

Dengue fever,  
Cardiac  
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### Introduction

Dengue Fever an acute febrile infectious disease, caused by any of four serotypes of a virus from the genus flavivirus, called dengue

virus. The highest incidence of dengue is seen in Southeast Asia, India and the American tropics (1-4). Dengue affects people of all ages. Severe dengue infections may give rise to many complications such as liver failure,

disseminated intravascular coagulation, encephalopathy, myocarditis, acute renal failure and haemolytic uraemic syndrome (5).

Although shock in DHF/DSS has been attributed largely to decreased intravascular volume due to capillary leakage of plasma into the interstitial space, a few recent studies have reported that it may be due to cardiac involvement. Acute reversible myocarditis has been reported in patients with dengue infections (6-9). ST segment and T wave changes in the electrocardiogram together with low ejection fraction and global hypokinesia on radionuclide ventriculography have been found. Thus Dengue fever is not only a serious illness it is emerging as a public health problem, so the need for the study.

In view of the increasing incidence of dengue fever as well as the rising morbidity and mortality due to this, the study of both the sub-clinical and clinical cardiac manifestations in dengue via clinical examination, cardiac biomarkers, ECG and ECHO cardiography was undertaken

The increase in capillary permeability that occurs in some patients, and can cause intravascular hypovolaemia and shock, is the best known cardiovascular complication associated with dengue.

Additionally, various specific cardiac manifestations have been described, ranging from rare fulminant myocarditis to more-common associations with functional myocardial impairment and arrhythmias (9-12). Myocarditis has now been included in the definition of severe dengue adopted in the 2009 WHO revised classification, but the true incidence of myocarditis remains unknown owing to the lack of screening in most countries where DENV is endemic (13-15).

In the past 2 decades, the critical role of myocardial impairment in the development of

septic shock has become clear, distinct from cardiovascular compromise caused by reduced preload and systemic vascular resistance. Myocardial impairment is possibly mediated by circulating myocardial depressant factors.

By contrast, the contribution of cardiac dysfunction to haemodynamic compromise in DSS remains to be adequately defined.

The main difficulty in describing the manifestations and frequency of cardiac involvement in dengue is the lack of clear criteria to define cardiac involvement.

Cardiac manifestations of dengue include functional myocardial impairment, arrhythmias, and myocarditis, which can occur through a number of mechanisms.

The wide clinical presentation of, and difficulties in diagnosing, myocarditis make the incidence difficult to quantify.

### **Aims and Objectives**

To determine the clinical profile of Dengue patients in the study and to Study the cardiac manifestations in patients with dengue fever.

### **Materials and Methods**

#### **Study site and study population**

The study is planned to be conducted in the Department of General Medicine, Karnataka Institute of Medical Sciences, Hubli, Karnataka

#### **Inclusion criteria**

Dengue fever cases aged more than 18 years attending at KIMS Hospital, Hubli.

Dengue fever cases confirmed by NS1Ag test or both NS1Ag and IgM, IgG positive cases.

### **Exclusion criteria**

Patients with previous history of any cardiac illness.

Admission ECG suggestive of old MI.

Medication affecting the heart rate.

Those who doesn't give consent for study

### **Study design**

This is a hospital based Observational study

### **Study period**

1 ½ year (1<sup>st</sup> March 2017 to August 2018)

### **Sample size**

Universal Sample size was taken in the study. I.e. all the cases of Dengue Fever admitted to KIMS Hospital, Hubli for the period of 1 ½ year.

### **Sampling method**

Purposive sampling method was used to recruit patients as and when admitted to Hospital.

### **Data collection**

This study was a hospital based study conducted in Karnataka Institute of Medical Sciences, Hubli. All the cases which satisfy the inclusion criteria were included in the study. Informed consent was taken and the clinical and laboratory parameters were recorded in predesigned data sheet. Diagnosis and management were done according to WHO guidelines. Serological tests for detection of IgM dengue antibodies and NS1 antigen was done. All the subjects were subjects for laboratory investigations.

The following investigations will be done in all case

Complete hemogram

Platelet count

Antigen test

Serology for dengue IgM, IgG, NS1 antigen

ECG

2D Echocardiography

### **Statistical analysis**

Data was entered into Microsoft excel data sheet and was analyzed using SPSS 22 version software. Categorical data was represented in the form of Frequencies and proportions. Chi-square test was used as test of significance for qualitative data. Continuous data was represented as mean and standard deviation.

### **Results and Discussion**

In the study majority 36% were in the age group 20 to 29 years, followed by <20 years (22%) and others as shown in table 1. Mean age of subjects was  $30.32 \pm 13.66$  years.

In the study 59% were males and 41% were females (Table 2).

In the study 100% presented with fever, 20% had associated headache, 4% had vomiting, 1% had chills, cough, giddiness and pain abdomen respectively and 2% had Myalgia. In the study 8% had Bradycardia and 20% had tachycardia. Mean Pulse rate was  $90.64 \pm 13.96$  bpm (Table 3).

In the study according to SBP, 6% had Hypotension and as per DBP, 6% had hypotension and 1% had hypertension.

Mean SBP was  $111.7 \pm 10.13$  mmhg and mean DBP was  $72.92 \pm 8.36$  mmhg (Table 4 and 5).

In the study 67% had no rashes, 24% had Petechial rashes, 4% had Ecchymosis (Table 6).

Petechie, 3% had Gum Bleeding, 1% had Bleeding in Oral cavity and Subconjunctival Hemorrhage respectively.

In the study 50% were NS1 positive, 20% were IgM positive, 12% were IgG Positive, 8% were IgM and IgG positive, 7% were IgM and NS1 Positive, 2% were NS1 and IgG Positive and 1% were IgG, IgM and NS1 Positive (Table 7).

In the study 31% had Hb <12 gm% and 69% had Hb >12 gm%. Mean Hb was  $12.68 \pm 2.12$  gm%. In the study 34% had Total count <4000 and 5% had Total count >11000. Mean Total count was  $5315.1 \pm 2771.24$  (Table 8).

In the study 26% had platelet count <20000, 32% had 20000 to 50000, 18% had 51000 to 100000 and 24% had platelet count >100000. Mean platelet count was  $58648 \pm 48496.45$  (Table 9 and 10).

In the study 18% were positive for Troponin I and 82% were negative for Troponin I. In the study 28% had CK-MB <25 and 72% had CK-MB >25. Mean CK-MB was  $85.99 \pm 75.73$  (Table 11 and 12).

**Table.1** Age distribution in the study

		Count	%
Age	<20 years	22	22.0%
	20 to 29 years	36	36.0%
	30 to 39 years	19	19.0%
	40 to 49 years	12	12.0%
	>50 years	11	11.0%
Total		100	100.0%

**Table.2** Sex distribution of subjects

		Count	%
Sex	Female	41	41.0%
	Male	59	59.0%
	Total	100	100.0%

**Table.3** Symptoms among subjects

		Count	%
Fever	Present	100	100.0%
Headache	Absent	40	40.0%
	Present	60	60.0%
Vomiting	Absent	94	94.0%
	Present	6	6.0%
Others	Chills	1	1.0%
	Cough	1	1.0%
	Giddiness	1	1.0%
	Myalgia	4	4.0%
	Nil	89	89.0%
	Pain Abdomen	4	4.0%

**Table.4** Pulse rate among subjects

		Count	%
Pulse Rate	Bradycardia	8	8.0%
	Normal Pulse Rate	72	72.0%
	Tachycardia	20	20.0%

**Table.5** Blood pressure among subjects

		Count	%
SBP	Hypotension	6	6.0%
	Normal BP	94	94.0%
DBP	Hypotension	6	6.0%
	Normal BP	93	93.0%
	Hypertension	1	1.0%

**Table.6** Rashes among subjects

		Count	%
Rashes+	Nil	67	67.0%
	Petechial rashes +	24	24.0%
	Ecchymosis,Petechie+	4	4.0%
	Gum Bleeding	3	3.0%
	Bleeding in Oral cavity	1	1.0%
	Subconjunctival Hemorrhage	1	1.0%

**Table.7** Dengue serology among subjects

		Count	%
Dengue Test	NS1 Positive	50	50.0%
	IgM positive	20	20.0%
	IgG Positive	12	12.0%
	IgM and IgG positive	8	8.0%
	IgM and NS1 Positive	7	7.0%
	NS1 and IgG Positive	2	2.0%
	IgG, IgM and NS1 Positive	1	1.0%
	Total	100	100.0%

**Table.8** Hb distribution among subjects

		Count	%
Hb	<12 gm%	31	31.0%
	>12 gm%	69	69.0%

**Table.9** Total count distribution among subjects

		Count	%
Total Count	<4000	34	34.0%
	4000 to 11000	61	61.0%
	>11000	5	5.0%

**Table.10** Platelet Count Distribution among subjects

		Count	%
Platelet count	<20000	26	26.0%
	20000 to 50000	32	32.0%
	51000 to 100000	18	18.0%
	>100000	24	24.0%

**Table.11** Troponin levels distribution among subjects

		Count	%
Troponin	Positive (>0.017)	18	18.0%
	Negative (<0.017)	82	82.0%
	Total	100	100.0%

**Table.12** CK-MB distribution among subjects

		Count	%
CK-MB	<25	12	28.0%
	>25	88	72.0%

**Table.13** ECG findings among subjects

		Count	%
ECG	Normal	56	56.0%
	Sinus Bradycardia	15	15.0%
	Sinus Tachycardia	9	9.0%
	T wave changes	10	10.0%
	ST Changes	3	3.0%
	Sinus Bradycardia with T Wave Changes	2	2.0%
	Sinus Tachycardia with T wave changes	2	2.0%
	LVH	1	1.0%
	Abnormal Rhythm	2	2%

**Table.14** ECHO findings among subjects

		Count	%
ECHO Findings	Normal	85	85.0%
	Anterior Wall Hypokinesia	6	6.0%
	Mild PAH and Mild TR	1	1.0%
	Mild PAH, Mild TR, Hypokinesia Ant.wall	4	4.0%
	LVH	1	1.0%
	DCM Secondary to Dengue Myocarditis	2	2.0%
	Mild AML Prolapse	1	1.0%

**Table.15** LFT findings among subjects

		Count	%
LFT	Abnormal	29	29.0%
	WNL	71	71.0%

**Table.16** Renal function tests in study subjects

	Number of Cases		Total
	Deranged RFT	Normal	
<b>Male</b>	5	54	59
<b>Female</b>	3	38	41
<b>Total</b>	8	92	100

In the study 56% had normal ECG, 15% had Sinus Bradycardia, 9% had Sinus Tachycardia, 10% had T wave changes, 3% had ST Changes, 2% had Sinus Bradycardia with T Wave Changes, 2% had Sinus Tachycardia with T wave changes and 1% had LVH and 2% had Abnormal Rhythm (Table 13).

In the study 85% had normal ECHO, 6% had Anterior Wall Hypokinesia, 1% had Mild PAH and Mild TR, Mild MR, Mild PAH, Mild TR, Hypokinesia Ant. wall, LVH, DCM Secondary. To dengue myocarditis and bradycardia, mild AML prolapse respectively (Table 14).

In the study 29% had abnormal Liver function tests. In the Study subjects: 5 Males and 3

Females had Deranged RFT and 54 Males and 38 Females had Normal RFT (Table 15 and 16).

Cardiac manifestations in dengue virus infection can range from asymptomatic Bradycardia to life threatening myocarditis (16-19). Various studies have quoted several cardiac manifestations of dengue infection viz. sinus Bradycardia, transient AV blocks, transient ventricular arrhythmias, myocarditis, systolic and diastolic dysfunction and pericardial effusion.

Myocardial involvement may be the direct result of dengue virus infection in susceptible individuals or may be due to effects of cytokines / cellular mediators of immune response.

The present study was done with the aim to determine the clinical profile of Dengue patients and to Study the cardiac manifestations in patients with dengue fever

### **Cardiac manifestations**

In the present study 56% had normal ECG and 44% had abnormal ECG findings, of them 15% had Sinus Bradycardia, 9% had Sinus Tachycardia, 10% had T wave changes, 3% had ST Changes, 2% had Sinus Bradycardia with T Wave Changes, 2% had Sinus Tachycardia with T wave changes and 1% had LVH and 2% had Abnormal Rhythm.

In the study by Kularatne, 62.5% of patients had abnormal ECG findings.

In the study by Sheethal *et al.*, the commonest rhythm abnormality noted was sinus bradycardia, found in 32%. Three patients had unexplained sinus tachycardia. In the study by Gupta *et al.*, sinus bradycardia was found in 14.28%, and sinus tachycardia 21.4 percent. AV dissociation with sinus node dysfunction was observed in one patient, which resolved in 24 hours. Kaushik *et al.*, have described atrioventricular dissociation and sino atrial exitblock in a child with dengue fever. Ventricular arrhythmias in the form of ventricular bigeminy, ventricular trigeminy and ventricular tachycardia was noted in one patient each. All these changes reverted back to sinus rhythm in 24 hours. Chuah *et al.*, and Veloso *et al.*, have described transient ventricular arrhythmias as a cardiac manifestation of dengue fever.

In the present study on ECHO, 6% had Anterior Wall Hypokinesia, 1% had Mild PAH and Mild TR, Mild MR, Mild PAH, Mild TR, Hypokinesia Anterior wall, LVH, 2% had DCM Secondary to Dengue Myocarditis And, Mild AML Prolapse Respectively.

In the study by Sheetal *et al.*, Echocardiographic evaluation was done in 18 patients. Three patients were noticed to have Mild pericardial effusion. Echocardiographic evidence of myocarditis was not seen in any patient. In the study by Gupta *et al.*, systolic dysfunction was absent in all patients; mild diastolic dysfunction was present in 14.28 percent. Wiwanitkit *et al.*, have described cases of dengue myocarditis.

In study by Carlos Henrique Miranda 2 patients had reduced LV function, two had LV segmental hypokinesia and one had pericardial effusion

The incidence of cardiac manifestations was present in 44% of the patients. A wide range of cardiac manifestations was observed in this study. CK-MB and Troponin I at admission were raised in 72% and 18% respectively. On ECG, 56% of the patients had normal rhythm, 15% had sinus Bradycardia, 9% had sinus tachycardia, 10% had T wave changes, 3% had ST Changes, 8% had Sinus Bradycardia with T Wave Changes, 2% had Sinus Tachycardia with T wave changes and 1% had LVH and 2% had Abnormal Rhythm. On 2D ECHO 6% had Anterior Wall Hypokinesia, 1% had LVH, and 2% DCM secondary to dengue myocarditis.

Hence Transient cardiac abnormality can be an important presentation among patients with Dengue fever and this should guide the treating physician to look for cardiac involvement

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