

# Evaluation of cardiac, hepatic enzyme levels and haematological profiles in chikungunya patients

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## Abstract

**Background:** Chikungunya is a viral infection caused by the Chikungunya virus (CHIKV) belonging to the Togaviridae family. The transmission of the virus is taking place through the bite of infected daytime biting female mosquitoes – primarily Aedes aegypti and Aedes albopictus. There were many reports which highlighted the number of reported cases of Chikungunya has been increasing in India in recent years, especially since 2016-2017.

**Materials and Methods:** Blood samples were collected from patients diagnosed with having Chikungunya. Cardiac profiles, routine laboratory investigations were carried out for analysing haematological and serum enzyme profiles.

**Result:** Our results highlighted that the alterations of cardiac profiles, haematological and hepatic dysfunction are prevalent in Chikungunya infection, with SGPT rising significantly more than SGOT. Hepatic Enzymes levels appear to have a directly proportional correlation with severity of infections.

**Conclusion:** Our study demonstrated that cardiac profiles, haematological and hepatic enzymes levels should be explored as routine laboratory markers for assessing the severity of Chikungunya infection, as they will help in employing an appropriate patient therapy, and thus optimise the use of available resources.

Keywords: cardiac profile, chikungunya, dengue, hepatic, haematological

## Introduction

Chikungunya affects all age groups, but severe complications are more often observed in children. Symptoms generally start 4-7 days after the mosquito bite. The acute phase is characterised by joint and muscle pain, high fever, extreme weakness, headache, vomiting, and rashes. In the chronic phase, various neurological syndromes and non-neurological manifestations can occur <sup>[1]</sup>.

Chikungunya (CHIK) is a mosquito-borne viral infection, characterised by a sudden onset of fever, severe joint and muscle pain that is sometimes accompanied by a skin rash. The symptoms, such as arthralgia, may last for months to years, especially in adults. In 2006, the CHIK virus (CHIKV) re-emerged in India after 32 years, causing the epidemic affecting more than 1.4 million people across the 13 States, and post epidemic, a declining trend was seen till 2011<sup>[1,3]</sup>.

The WHO South-East Asia Regional Office has reported a significant number of Chikungunya cases in almost 151 districts in 8 states of India suffered from many cases of chikungunya fever. The most affected states are Andhra Pradesh, Andaman & Nicobar Islands, Tamil Nadu, Karnataka, Maharashtra, Gujarat, Madhya Pradesh, Kerala and Delhi <sup>[1, 2]</sup>.

As usual, the outbreaks of Chikungunya infections are most likely to occur in the post-monsoon period when the vector density significantly increases. Human beings are considered as the reservoir for the chikungunya virus during the epidemic time.

The rapid spread of the disease in Southern part of India from 2004 has afflicted millions of people and left many of them with crippling disabilities. Chikungunya disease continues to cause epidemics in many countries in the region. The patient in this context acts as the reservoir of infection for others in the household and the community. Therefore, public health measures to minimise the transmission of infection become imperative to prevent and control the outbreak from spreading <sup>[3, 5]</sup>.

Fever with or without arthralgia is a prevalent manifestation of several other diseases. CHIK fever may not have typical manifestations, or it may co-exist with other infectious diseases like dengue fever or non-infectious diseases like rheumatoid arthritis<sup>[6]</sup>.

Three stages of disease after the incubation period in CHIK have been recognized as Acute, sub-acute and chronic. Not every infected individual develops the full three stages it is also been reported that severe acute manifestations of the disease, especially in which the liver, lungs, and even the eye are affected by the extra-articular intense inflammatory response. Similarly, the involvement of the heart has often been fatal and worth highlighting in some reports, but it has not been very largely discussed. But, the mechanism of cardiac involvement in Chikungunya fever is not fully understood <sup>[7]</sup>.

The analysis and diagnosis of the case in the community need to be communicated immediately to the nearest public health official for identification of clusters by person, place and time and expansion of the control measures in the community and district levels.

In this study we tried to establish the relationship between the severity of Chikungunya infection and its impact on Levels hepatic enzymes and haematological profiles of individuals.

#### **Materials and Methods**

Study Design: It was a prospective cohort study over two years through sample and sampling techniques. A total of 50

patients admitted to the hospital with a history of fever of more than 38.50C and CHICKV positive were selected using purposive sampling techniques. Clinical signs and symptoms, severity, and outcomes with relevant laboratory parameters were compared in detail. They are followed from the onset of fever to time of recovery or discharge according to WHO discharge criteria whichever is earlier. Same numbers of normal healthy individuals were also recruited for this study. Duly signed Informed consent forms were collected from all the participants during the study.

Routine Laboratory Investigations: The 5-7ml of blood samples were collected from all these participants' investigations were done- blood counts, clinically patients are monitored and for platelets, hematocrit count, haemoglobin count, total leukocyte count.

Liver Function Test: SGPT, SGOT were estimated by using protocol prescribed by the International Federation of Clinical Chemistry. ALP, Total Protein, Albumin, Total Bilirubin were estimated by colourimetric assay.

Cardiac studies included, the main clinical features, ECG's, chest radiographs, virological investigations were performed.

Statistical analysis: The collected data analysis was done using SPSS software. The results were presented as mean  $\pm$ standard deviation (SD) and percentages. Mean, and SD was computed for all continuous variables and comparisons were done using Student's t-test.

#### Results

The total numbers of 50 cases of acute febrile illness were recruited based on their clinical report as seropositive for CHIKV infection, an equal number of healthy individuals also participated in this study. There were altogether 67 (63.6%) men and 33 (36.4%) women participants. Most of the participants were in the 20–30-year age group. A seasonal peak with 42 (34.7%) cases was seen in monsoon season. The patient characteristics and salient clinical features of acute chikungunya infection, and biochemical parameters are compared in Tables 1 and 2. The rural residency was prominent in all groups. Fever was seen in all cases (100%). The symptomology of chikungunya included headaches, joint pain, weakness, and restricted joint movement. No complications were reported. Deranged liver function was seen in all patients (100%).

**Table 1:** Comparison of characteristics and clinical features in between normal healthy individuals and patients groups.

Characteristics		<b>Control Group</b>	Patient Group
Age Group (Years)	<20	4 (2)	4 (8)
	20-30	21 (42)	27 (54)
	31-40	12 (24)	11 (22)
	41-50	8 (16)	3 (6)
	51-60	5 (10)	2 (4)
	>60	0	3 (6)
Gender	Male	40 (80)	37 (74)
	Female	10 (20)	23 (46)
Residence	Urban	24 (48)	22 (44)
	Rural	26 (52)	28 (56)
Symptoms	Fever	2 (4)	50 (100)
	Joint Pain	1 (2)	31 (62)
	Backache	0	15 (30)
	Vomiting	0	4 (8)

Out of 50 patients, 37 were male and 23 were female and in control group of 50 healthy individuals 40 were male and 10 were female. Most of the participants were in the age group of 20 - 40 and none of them were below 20 years and above 70 years of age. All the patients had fever as a presenting complaint. Most of them had myalgia at the time of presentation (Table 1).

 
 Table 2: Comparison of biochemical parameters in between normal healthy individuals and patients groups.

Parameters	<b>Control Group</b>	<b>Patient Group</b>	p Value
Hb	13.15±1.82	11.55±2.36	0.5012
TLC	8560±2634	4781±2219	0.0082
PC	155,700±23,523	67,622±44,254	0.0015
HCT	36.67±4.52	38.66±6.07	0.523
SGPT	28.87±4.23	239.68±589.25	0.5215
SGOT	$25.92 \pm 6.22$	$345.52 \pm 1356.52$	0.633
ALP	34.21±14.21	144.21±110.26	0.2231
Total Bilirubin	0.98±0.29	$1.84 \pm 0.28$	0.001
Direct Bilirubin	0.51±0.18	0.98±0.33	0.001
Total Protein	7.4±0.59	6.29±0.89	0.0747
Albumin	4.11±0.47	3.15±0.69	0.0130

Hb: Hemoglobin, TLC: Total leukocyte count, PC: Platelets count, HCT: Hematocrit, RBS: Random blood sugar, ALP: Serum alkaline phosphatase, SGPT: Serum glutamate pyruvate transaminase, SGOT: Serum glutamic-oxaloacetic transaminase.

Elevated liver enzymes were found in almost all patients. Of the liver enzymes, SGPT levels were significantly higher than SGOT levels.

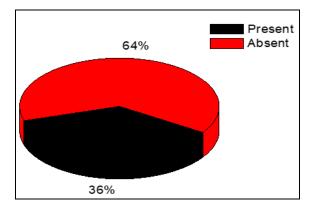


Fig 1: Distribution of individuals according to Cardiac manifestations

The cardiac manifestations studied in the present showed, out of 50 patients, 64 % were identified with cardiac manifestations, whereas, 18% of the patients were absent for cardiac manifestations (Figure 1).

Table 3: Comparison of deranged liver enzymes in between
normal healthy individuals and patients groups.

Para	meters	<b>Control Group</b>	<b>Patient Group</b>
SGPT (U/L)	60-180 (%)	1 (2)	29 (58)
	181-500 (%)	0	12 (24)
	501-1000 (%)	0	5 (10)
	>1000 (%)	0	4 (8)
SGOT (U/L)	38-120 (%)	0	25 (50)
	121-500 (%)	1 (2)	13 (26)
	500-1000 (%)	0	9 (18)
	>1000 (%)	0	3 (6)

 
 Table 4: Comparison of thrombocytopenia in between normal healthy individuals and patients groups.

Platelets Count (/µL)	<b>Control Group</b>	<b>Patient Group</b>
>150,000	26 (52)	2 (4)
150,000-100,000	23 (46)	5 (10)
100,000-50,000	1(2)	20 (40)
50,000-20,000	0	13 (26)
<20,000	0	6 (12)
<10,000	0	3 (6)
Platelets transfusion	0	1 (2)

Thrombocytopenia was noted in 43 patients, of whom 12 had bleeding, and 31 had no bleeding. Statistically, the Chisquare test was used to test the significance of thrombocytopenia and bleeding tendencies. There was no significant difference (p = 0.945) found between the variations in the thrombocytopenia and the bleeding tendencies. This implies that bleeding tendencies does not depend upon thrombocytopenia.

## Discussion

Currently, Chikungunya is causing significant public health concern throughout the world, particularly in South-East Asian countries. Recently dengue outbreaks caused significant morbidity and mortality in certain parts of India mainly in Andhra Pradesh, Uttar Pradesh and Tamil Nadu <sup>[8]</sup>. Hepatic dysfunctions in dengue are common. It is due to either direct effect of the virus on hepatocytes or due to reactive hepatitis. Hepatic involvement in dengue fever is in the form of elevated serum aminotransferase [9]. Those patients with elevated liver enzymes are more likely to have increased risk of bleeding tendencies, shock, ARDS, renal failure. In addition to decreased platelet count, hepatic dysfunction plays a significant role in bleeding. Hence, it is mandatory to evaluate serum aminotransferases in all patients with dengue fever <sup>[10]</sup>. It is observed that, out of 50 patients, most of the patients had elevated liver enzymes, SGPT levels were significantly higher than SGOT levels, and these patients had severe complications.

Chikungunya viral infection is not generally fatal but can cause neurological and optical manifestations. Severe joint pain is the prominent clinical manifestation and can persist for months to a year <sup>[11]</sup>.

As Dengue Fever (DF) has a high incidence and mortality rate, symptomatic patients are tested only for DENV and only in rare cases for chikungunya viral infection. This scenario is an important reason why chikungunya cases go undiagnosed in dengue-endemic regions, and the actual burden of the chikungunya viral infection has been missed. Thus, investigation for both viruses should be done, especially in endemic regions. Accurate and early diagnosis of coinfections would help inappropriate management <sup>[12, 13]</sup>.

The number of cases increases during and after the monsoon months because higher humidity lengthens the life span of mosquitoes and increased temperatures shorten the extrinsic incubation period <sup>[14]</sup>. Rising of the cases during the monsoon period with highest cases in different months had been reported in previous studies from India <sup>[12, 15]</sup>. November was observed as a peak among all the months of the year in this study.

Earlier studies have reported increased incidences of atypical hemorrhagic manifestations <sup>[13, 15]</sup> and hepatic dysfunction <sup>[16]</sup>. Hepatic dysfunction was the most common

complication in our dengue and coinfection groups. SGPT levels were significantly higher than SGOT levels.

## Conclusion

In Chikungunya cardiac profiles, Haematological profile alteration and Hepatic involvement are common. It is characterized by elevated liver enzymes, i.e. SGPT more than SGOT levels. Elevated liver enzymes are associated with complications like bleeding, shock and organ impairment. In addition to thrombocytopenia, hepatic involvement plays a significant role in bleeding. Elevated liver enzymes have got prognostic value in this study. Hence, liver enzymes are mandatory in dengue fever to look for complications, and it is of prognostic value. Those patients with elevated liver enzymes should be monitored carefully than those patients with normal liver enzymes.

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