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Clinical Profile and Laboratory Findings of Dengue Fever in Children

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ABSTRACT

Background: Dengue is an arthropod borne viral hemorrhagic fever. It is a major public health concern throughout the tropical and subtropical regions of the world.

Objectives: To study clinical profile and laboratory findings of dengue infection in children.

Materials and Methods: Children up to the age of 15 years who were confirmed serologically dengue positive (either positive NS1 antigen or positive IgM antibody by ELISA method were included in study. Total 100 patients were studied. Pre tested, semi structured questionnaire were developed and used for data collection.

Result and Discussion: In current study, maximum patients were found in Dengue Hemorrhagic Fever (DHF) group (68%), followed by Dengue Fever (25%). Fever was most common clinical manifestation. Maximum patients in this study were in 10 to 15 years of age group, male being more commonly affected. Thrombocytopenia and leukopenia were present in 90% and 69% of patients respectively.

Conclusion: School going children were more exposed to mosquito bites during day time school activities, thus maximum patients were from 10 to 15 years of age group. In early diagnosis of dengue fever, ELISA NS1 antigen is seen more promisable. When used in combination with ELISA IgM antibody, it significantly improves the diagnostic algorithm. Thrombocytopenia is more consistent feature of Dengue Hemorrhagic Fever (DHF) and Dengue Shock Syndrome (DSS).

Keywords: Dengue Hemorrhagic Fever (DHF), Dengue Shock Syndrome (DSS), ELISA NS1 antigen, ELISA IgM antibody

INTRODUCTION

Dengue is the most rapidly spreading mosquito-borne viral disease of mankind, with a 30- fold increase in global incidence over the last three decades¹. Dengue fever is an acute febrile infectious disease in subtropical and tropical areas that is progressively making its way from being "one of the great neglected diseases of mankind" towards being acknowledged as one of the world's major infectious

disease^{2,3}. Dengue is a fast emerging and rapidly spreading systemic viral infection with global estimates of 390 million infections per year, of which 96 million are apparent infections and 3.97 billion people in 128 countries are at risk of dengue infection⁴. Dengue is one of the most common vector-borne diseases in Southeast Asia and has been ranked as the most important mosquito-borne viral disease with epidemic potential in the world⁵. Some 2.5

billion people - two fifths of the world's population in tropical and subtropical countries are at risk⁶. The etiologic agents include all four serotypes which belong to genus Flavivirus in the family Flaviviridae⁷. Four serotypes, dengue known as DENV-1, DENV-2, DENV-3, and DENV-4, are genetically related and biological similar, can cause severe and fatal disease⁸. The principle vector is female mosquito, Aedes aegypti, which breeds largely indoors in clean waters mainly in artificial water containers, and feeds on daytime⁹. humans in The manifestations of dengue vary with the age and immunity of the patient. It can present as 1) an inapparent infection 2) nonspecific febrile illness, 3) classic dengue fever, 4) dengue hemorrhagic fever, 5) Dengue shock syndrome (DSS) and 6) encephalopathy and 7) fulminant liver Failure¹⁰.

After incubation, the disease begins abruptly and & clinical course of illness passes through the following three phases:

- 1) Febrile phase
- 2) Critical phase
- 3) Convalescent phase^{11,12}. The present study was carried out to study clinical profile and laboratory findings, of dengue infection in city and rural area.

AIMS AND OBJECTIVES

- To study the clinical profile of patients with dengue fever.
- To study the various laboratory parameters in patients with dengue fever.

MATERIALS AND METHODOLOGY STUDY DESIGN:-

A prospective observational study.

STUDY DURATION:-

Study Duration was 18 months, from 1st January 2018 till 30th June 2019.

STUDY POPULATION:-

The present study was conducted on children up to 15 years of age who were

confirmed serological dengue positive (either Positive NS1 antigen or Positive Ig M antibody by ELISA method), referred to tertiary care hospital.

SAMPLE SIZE:-

Sample size was 100 children up to 15 years of age who fulfilled the inclusion criteria.

SAMPLING TECHNIQUE (SAMPLING METHOD):-

Simple random sampling technique was used.

SELECTION CRITERIA:-INCLUSION CRITERIA:-

- 1. All patients between the age group of 0 15 years.
- 2. Patients with confirmed diagnosis of dengue fever (positive test for NS1 antigen/IgM antibody by ELISA method).

EXCLUSION CRITERIA:-

1. Patient with serological test negative for dengue.

METHODOLOGY OF STUDY:

The pre-tested, semi structured questionnaire developed and used for data collection. Patient / Patient guardian/ Patient relative were informed and explained about the study. Written informed consent had been taken prior to procedures as per attached proforma.

- Socioeconomic data was collected.
- Clinical histories were noted down along with thorough clinical examination.
- Other co- morbid condition and complications were noted down.
- Relevant laboratory investigations were carried out.
- The following laboratory investigations were carried out as per clinical management protocol.
- Hb
- PCV
- WBC
- Platelets

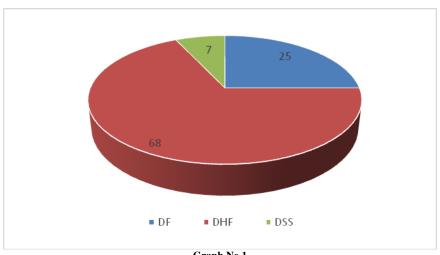
- NS1 antigen
- IgM Dengue
- IgG Dengue
- LFT
- RFT
- PT INR
- X- ray chest
- Clinical and consequential data and interpretation laboratory were documented by a predesigned proforma.
- Results were judged, based on the clinical examinations, observations, laboratory investigation and radiological interpretation as per proforma.
- The observations were graphically depicted and with the help of tables and

- conclusions are drawn on the basis of observations and discussion.
- The data obtained were compiled in tabular form and analysed. Descriptive statistics were used to present data in tabular and graphical forms and suitable statistical tests applied for results.

OBSERVATION AND RESULTS

1) Distribution of cases according to the diagnosis.

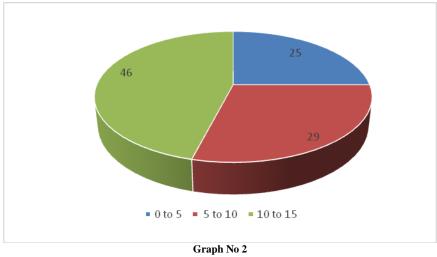
In the current study, maximum patients were found in the DHF group (68 %) followed by DF (25 %) and DSS (7 %) respectively.



Graph No 1

2) Age and gender wise distribution of cases

Maximum patients in the current study were in the 10 to 15 year age group (46%), followed by 5-10 years of age group (29%). Males are affected more than the females.



3) Clinical manifestations

Table no 1:

| Clinical manifestations | Number of cases | Percentage | |
|----------------------------|-----------------|------------|--|
| Fever | 100 | 100% | |
| Headache | 26 | 26% | |
| Retro orbital pain | 17 | 17% | |
| Arthralgia | 18 | 18% | |
| Body ache | 25 | 25% | |
| Nausea and vomiting | 52 | 52% | |
| Diarrhea | 9 | 9% | |
| Abdominal pain | 25 | 25% | |
| Rash | 54 | 54% | |
| Conjunctival congestion | 13 | 13% | |
| Hemorrhagic manifestations | 58 | 58% | |

The above table shows that in the current study, fever is the most common clinical feature in dengue (100%) followed by Hemorrhagic manifestations (58%).

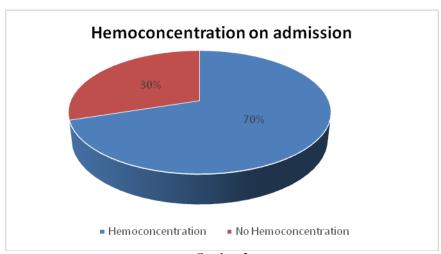
4) Elisa NS 1 antigen test, IgM Antibody Test, IgG dengue test

ELISANS 1 antigen test was done in all patients out of which 70 were positive. In 30 patients NS 1 antigen test was negative.

ELISA IgM antibody test was done in all patients, out of which 37 were positive. Both Elisa Ns1 antigen and IgM antibody were positive in 9 patients. IgG dengue test was positive in 2 patients.

5) Hemoconcentration on admission

Hemoconcentration was present in 70% patients on admission



Graph no 3

6) On admission, thrombocytopenia:

Table no 3:

| Thrombocytopenia (Platelet < 1.5 L/Cu mm | | | | | |
|--|---|--------|--------|--------|--------|
| Yes | N | DF | DHF | DSS | Total |
| | % | 22 | 61 | 7 | 90 |
| No | N | 24.4% | 67.8% | 7.8% | 100.0% |
| | % | 3 | 7 | 0 | 10 |
| Total | N | 30.0% | 70.0% | 0.0% | 100.0% |
| | % | 25 | 68 | 7 | 100 |
| | | 100.0% | 100.0% | 100.0% | 100.0% |

The above table shows that thrombocytopenia was present in 100% of the patients.

7) On admission, leukopenia

Leukopenia was present in 69% of patients

DISCUSSION

1) Distribution Of Cases According To Diagnosis

In the current study, 25% of the patients were of DF, 68% were of DHF and 7% were of DSS. In study conducted by Manjith Narayan et al (2000), 33.8% of patients were of DF, 57.6% were of DHF and 8.47% were of DSS¹³. In the study by Hema Mittal et al (2010), 8% of the patients were of DF, 51% were of DHF and 42% DSS¹⁴. In a study conducted by Ashwini Kumar et al (2010), out of total patients admitted, 83.9% had DF, 8.8% had

DHF and 7.3% had DSS¹⁵. We got maximum patients with DHF and DSS as our institute is tertiary referral care centre.

Table No 1

| Diagnosis | In current study | Manjith Narayan et al (2000) | Hema Mittal et al (2010) | Ashwin Kumar et al (2010) |
|-----------|------------------------|------------------------------------|--------------------------------|---------------------------------|
| DF | 25% | 33.8% | 8% | 83.9% |
| DHF | 68% | 57.6% | 51% | 8.8% |
| DSS | 7% | 8.47% | 42% | 7.3% |

2) Distribution of Cases According to Age Group.

In the current study, maximum patients were in the 10 to 15 year age group (46%), followed by 5 to 10 year age group (29%). In the study by **Senthil Kumar K et al(2018)**, maximum patients were in the more than 10 years age group (50.7 %)¹⁶. In a study conducted by **Shubhankar Mishra et al(2010)** maximum patients were in the age group more than 11 years(34 %)¹⁷. This shows that dengue affects school going children more. More prevalence of this age group is probably due to more involvement in outdoor activities during day time exposing them to higher risk of mosquito bites.

3) Male: Female Ratio Among Different Studies

In the current study, male: female ratio was 2.25:1, as compared to 1.32:1 in the study by **Mittal Hema et al (2010)**¹⁴ and 3.40:1 in the study by **Shubhankar et al (2016)**¹⁷. In a study conducted by **Ashwini Kumar et al (2010)**, male: female ratio was 1.83:1¹⁵.

4) Clinical features among different studies

In all studies, fever was the most common clinical feature (100%). In the current study, the other common features were hemorrhagic manifestations (58%), rash(54%), nausea/ vomiting (52%), respiratory symptoms (i.e. cough, cold) (35%), bodyache (25%), abdominal pain (22%). Other manifestations are conjunctival suffusion(13%) and diarrhoea (9%).

Fever was invariably high grade with temperature more than 39°C. Fever in dengue was not responding to normal dose of paracetamol.

Rash was more commonly seen on arms secondary to BPmonitoring (Torniquet test). When there is increased tendency, bleeding spontaneous ecchymosis, easy bruising and bleeding at the site of venepuncture were common. Some patients had gross ecchymosis or gastrointestinal bleeding, usually after period of uncorrected shock.

In the study by **Ashwin Kumar et al** (2010), the common features apart from fever were myalgia (64.6%) and vomiting (47.6%), headache (47.6%) and abdominal pain (37.6%). In the study by **Ragini Singh et al** (2013), the common features apart from fever were headache (80%) and myalgia (73.6%)²⁰. Rash was present in 15% cases. In the study by **Mittal Hema et al** (2002)¹⁴, bleeding manifestations were found in 48.8% of the cases. In a study conducted by **Senthil Kumar K** (2018), fever was present in 93 % patient. Other features were vomiting (68 %), headache (36%)¹⁶.

Table no 2:

| Table no 2. | | | | | | |
|----------------------------|-----------------|--------------------------|----------------------------|---------------------------|---------------------------------|--|
| Clinical Features | In Our Study | Hema Mittal et al (2002) | Ashwini Kumar et al (2010) | Ragini Singh et al (2013) | Senthil kumar K et al (2018) | |
| Fever | 100% | 100% | 99.1% | 100% | 93% | |
| Headache | 26% | 63% | 47.6% | 80% | 36% | |
| Retroorbital Pain | 17% | - | - | - | - | |
| Arthralgia | 18% | - | - | - | - | |
| Body ache | 25% | - | 64.6% | 73.6% | 15% | |
| Nausea and Vomiting | 52% | - | 47.6% | 11.4% | 68% | |
| Diarrhoea | 9% | - | 13.9% | 9% | 1.2% | |
| Abdominal Pain | 25% | 71% | 37.6% | 13.6% | 20% | |
| Rash | 54% | 26.6% | 21.7% | 15% | 20% | |
| Hemorrhagic Manifestations | 58% | 48.8% | - | 7.1% | 9% | |
| Conjunctival Congestion | 13% | - | - | = | - | |

5) Elisa NS1 /Igm /IgG

In the current study, Elisa NS 1 antigen test was done in all patients out of which 68% were positive. ELISA IgM antibody Dengue was found to be positive in 30% patients and IgG dengue was positive in 2% patients. Both Elisa NS1 and IgM were positive in 9% patients. In a study conducted by Senthil Kumar et al (2018), Elisa NS1 antigen assay was positive in 79.1% patients and Elisa Igm antibody was positive in 14.9% patients¹⁶.

In our study, majority of patients were positive for Elisa NS1 followed by Elisa IgM as large number of patients were present within day of illness 5. Elisa Ns1 antigen test helps in early detection of cases.

6) Haematocrit studies

In the study, current haemoconcentration was on admission present 63% of the in cases. Haemoconcentration on admission was one of the prominent features in the current study. This was due to the fact that many patients presented late in the course of the illness.

Usually on days 3-7 of illness, an increase in capillary permeability in parallel with increasing haematocrit levels may occur. This marks the beginning of the critical phase. The degree of increase above the baseline hematocrit often reflects the severity of plasma leakage. A hematocrit level increase greater than 20% is a sign of hemoconcentration and precedes shock. Thus, serial monitoring of haematocrit levels helps in management of patients with dengue.

In the study by **Narayan M et al** (1998), haemoconcentration was present in 40% of the cases.

7) Leukocyte counts

In the current study, leukopenia on admission was present in 69% of the total dengue patients. There was significant statistical difference among the three groups. In the study by **Ch Manoj Kumar et al (2018)**, leukopenia was present in

37.5% of the cases¹⁸ and it was 41.4% in the study by **Ragini Singh et al (2013)**²⁰ and 38.9% in the study by **Senthil Kumar K et al (2018)**¹⁶. Thus leukopenia on admission was one of the prominent features in the current study.

8) Platelet counts

In the current study, thrombocytopenia on admission was present in 90% of the total dengue patients. In study conducted by **Mahesh Ahirrao et al (2019)** platelet count was less than 100000 in 51 % patients¹⁹. In the study by **Mittal Hema et al (2002)**, thrombocytopenia was present in 92.6% of the cases¹⁴. In the current study, thrombocytopenia was found more commonly in DHF and DSS.

Platelet counts less than 100,000 cells/ μL are seen in dengue hemorrhagic fever or dengue shock syndrome and occur before defervescence and the onset of shock. Thus, the platelet count monitoring at least every 24 hours helps in early recognition of DHF and DSS.

CONCLUSION

- Maximum patients in our study were from 10-15 years of age group, school going children are more exposed to mosquito bites during day time school activities.
- 2) High grade fever, not responding to routine antipyretics dose should arouse a suspicion of Dengue fever in an epidemic setting.
- 3) Any fever associated with headache, body ache, erythematous moribiliform rash and conjuctival suffusion should be subjected to Dengue antigen test.
- 4) ELISA NS1 Ag assay holds promise in early diagnosis of dengue infection. When used in combination with ELISA IgM antibody, it significantly improves the diagnostic algorithm. Early identification of dengue infection using ELISA NS1 antigen test is valuable in terms of disease progression and mortality.

- 5) Any patient with symptoms of high grade fever and leukopenia on CBC should be investigated for Dengue fever.
- 6) Thrombocytopenia is most consistent feature of Dengue hemorrhagic fever and Dengue Shock Syndrome. Many patients with mild dengue fever may have normal platelet count.

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Ethical Approval: Approved

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