A STUDY ON CLINICAL PROFILE AND OUTCOME OF DENGUE FEVER IN CHILDREN ADMITTED IN A TERTIARY CARE HOSPITAL

Dr. Anitha Edara¹, Dr. M. Hema Sundar Rao², *Dr. Dileep Kumar Allagadda³

 Assistant Professor, Department of Pediatrics, Santhiram Medical College and Hospital, Nandyal, Andhra Pradesh.
 Associate Professor, Department of Pediatrics, Gayatri vidya Parishad Health care and medical technology, Visakhapatnam, Andhra Pradesh.
 Assistant Professor, Department of Pediatrics, Mallareddy Institute of Medical Sciences, Hyderabad.
 *Corresponding Author: Dr. Dileep Kumar Allagadda, Assistant Professor, Department of Pediatrics, Mallareddy Institute of Medical Sciences, Hyderabad.

ABSTRACT

Background:Dengue infection is the most rapidly emerging vector-borne viral disease with a 30-fold increasing global incidence over the last five decades. It is a major public health concern throughout tropical and subtropical regions of the world. In India, every year cases are spreading to newer geographical areas.

Objective: To study the clinical profile and outcome of Dengue fever in children admitted in a tertiary care hospital.

MATERIAL & METHODS: Study Design: Prospective Observational study.**Study area:** The study was done in Department of paediatrics, in a tertiary care hospital. **Study Period:**April 2021 – March 2022 (1 year).**Study population:**Children age group of 1-12 years presenting with fever and other features suggestive of Dengue fever.**Sample size:** A total of 62 were included in the study.**Ethical consideration:** Institutional Ethical committee permission was taken prior to the commencement of the study. **Study tools and Data collection procedure:**Children age group of 1-12 years presenting with fever and other features suggestive of Dengue fever according to WHO guidelines will be assessed clinically, serologically and managed as for WHO protocol and will be followed for outcome.

Results: Major clinical features that were observed in most cases were pain abdomen (74%), vomiting (72.5%), arthralgia (74%), body pains (72.5%), poor intake (74%), conjunctival suffusion (74%) & facial puffiness (74%), hepatomegaly (74%) and ascites (72.5%).

CONCLUSION:

The detection of IgM dengue antibodies by capture ELISA & NS1 Ag were helpful for diagnosis of acute dengue virus infection. The serological diagnosis of dengue fever has a role in categorizing primary and

secondary infection and it also serves as a predictor of disease progression and mortality especially in severe dengue.

Key words: IgM dengue antibodies, Dengue infection, ELISA & NS1 Ag

INTRODUCTION:

Dengue infection is the most rapidly emerging vector-borne viral disease with a 30-fold increasing global incidence over the last five decades. It is a major public health concern throughout tropical and subtropical regions of the world. In India, every year cases are spreading to newer geographical areas.

Dengue infection needs to be addressed as a single disease with different clinical presentations ranging from asymptomatic conditions to severe clinical courses that may lead to high morbidity and mortality. In the absence of a specific antiviral drug for dengue infection, it is a great challenge for the clinicians to recognize the severity of the disease at the early phase for early intervention and timely effective management to reduce complication and death¹.

The incidence of dengue has grown dramatically around the world in recent decades. The actual numbers of dengue cases are underreported and many cases are misclassified. One recent estimate indicates 390 million dengue infections per year (95% credible interval 284–528 million), of which 96 million (67–136 million) manifest clinically (with any severity of disease)¹. Another study, of the prevalence of dengue, estimates that 3.9 billion people, in 128 countries, are at risk of infection with dengue viruses². Member States in three WHO regions regularly report theannual number of cases across the Americas, South-East Asia and Western Pacific exceeded 1.2 million in 2008 and over 3.2 million in 2015².

The year 2016 was characterized by large dengue outbreaks worldwide. The Region of the Americas region reported more than 2.38 million cases in 2016, where Brazil alone contributed slightly less than 1.5 million cases, approximately three times higher than in 2014^2 .

In India, dengue has seen resurgence in recent times. The first evidence of Dengue fever in India was reported during 1956 in Vellore district (Tamil Nadu). The first Dengue hemorrhagic fever outbreak occurred in Calcutta (West Bengal) in 1963 with 30% of cases showing hemorrhagic manifestations. All of the four serotypes i.e. Dengue 1, 2, 3 and 4 have been isolated in India. Aedes aegypti breeds more commonly in urban areas. However the trend is now changing due to socioeconomic and manmade ecological changes, it has resulted in invasion of Aedes aegypti in to the rural areas, which has tremendously increased the chances of spread of the disease to rural areas. Recurring outbreaks of DF & DHF have been reported from various states/union

territories namely Andhra Pradesh, Delhi, Goa, Haryana, Gujarat, Karnataka, Kerala, Maharashtra, Rajasthan, Uttar Pradesh, Pondicherry, Punjab, Tamil Nadu, West Bengal and Chandigarh⁴. In 2015, Delhi, India, recorded its worst outbreak since 2006 with over 15000 cases ³.

Objective: To study the clinical profile and outcome of Dengue fever in children admittedin a tertiary care hospital.

MATERIAL & METHODS:

Study Design: Prospective Observational study.

Study area: The study was done in Department of paediatrics, In a tertiary care Hospital.

Study Period: April 2021 – March 2022 (1 year).

Study population:Children age group of 1-12 years presenting with fever and other features suggestive of Dengue fever.

Sample size: A total of 62 were included in the study.

Sampling method: Simple random sampling technique.

Inclusion Criteria:

• Children age group 1 - 12 years.

• Children's with fever and other features suggestive dengue fever according to WHO guidelines {headache,

retro orbital pain, myalgia / arthralgia, rash, hemorrhagic manifestations, thrombocytopenia and leucopenia }.

EXCLUSION CRITERIA:

- Those with other viral fevers with thrombocytopenia.
- Those with positive for Malaria parasite (All species).
- Those with acute and chronic liver disease.
- Those with blood dyscrasias.

Ethical consideration: Institutional Ethical committee permission was taken prior to the commencement of the study.

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Study tools and Data collection procedure:Children age group of 1-12 years presenting with fever and other features suggestive of Dengue fever according to WHO guidelines will be assessed clinically, serologically and

SEX	NO OF PATIENTS	PERCENTAGE
MALE	35	56.5
FEMALE	27	43.5
TOTAL	62	100

managed as for WHO protocol and will be followed for outcome.

All the children are subjected for following investigations

- Complete Blood Picture.
- IgM antibody detection (SD Dengue IgM Capture Elisa kit).
- NS1Antigen detection (Panbio Dengue Early ELISA kit).

Other relevant investigations for renal, liver and other functions.

The Panbio Dengue Early ELISA is a dengue NS1 antigen capture ELISA. It is for qualitative detection of NS1

Ag in human serum. SD Dengue IgM Capture Elisa kit is used for qualitative detection of IgM dengue

antibodies specific to Dengue virus in human serum.

OBSERVATIONS & RESULTS

Table 1: DISTRIBUTION OF CHILDREN ACCORDING TO GENDER

In this study 35(56.5%) were male children and remaining 27(43.5%) were female children.

Table 2: THE DISTRIBUTION OF DENGUE FEVER ACCORDING TO AGE

AGE GROUP IN YEARS	NO OF PATIENTS	PERCENTAGE
1-3	18	29
4-6	25	40.3
7-9	11	17.7
10-12	8	13
T0TAL	62	100

The study observed that maximum number of cases occurred between 4-6 yrs, constituting 40.3%.

Table 3: CLINICAL FEAUTRES

CINICAL FEAUTRES		NO OF PATIENTS		PERCENTAGE
HEADACHE Clinical picture	NO	OF PATIENTS	PE	REENTAGE
RETROORBITAL PAIN		21		33.8
FATIUGE		32		51.6
PAIN ABDOMEN		46		74.1
VOMITINGS		45		72.5
ARTHRALGIA		46		74.1
BODY PAINS		45		72.5
POOR INTAKE		46		74.1
SKIN BLEEDS		29		46.7
EPISTAXIS		20		32.2
HEMATEMESIS		13		20.9
MELENA		26		41.9
CONVULSIONS		2		3.2
CONJUCTIVAL SUFFUSION		46		74.1
HEPATOMEGALY		46		74.1
SPLENOMEGALY		32		516
TORNIQUET TEST		32		51.6
FACIAL PUFFINESS		46		74.1
ASCITES		45		72.5
PEDAL EDEMA		20		32.2

Major clinical features that were observed in most cases were pain abdomen (74%), vomiting (72.5%), arthralgia (74%), body pains (72.5%), poor intake (74%), conjunctival suffusion (74%) & facial puffiness (74%), hepatomegaly (74%) and ascites (72.5%).

Table 4: Classification of dengue according to clinical picture

DW	IgM Ab categorization		total(%)
	Positive (%)	Negative (%)	
Positive	24 (99.6)	1 (0.4)	25 (100)
Negative	6 (16.2)	31 (83.8)	37(100)
total	30 (48.3)	32(51.7)	62
- S NO			
1	Probable dengue	8	13
2	Dengue without warning signs	3	4.8
3	Dengue with warning signs	25	40.3
4	Severe dengue	26	41.9

Majority patients observed in this study were severe dengue (41.9%) followed by dengue with warning signs (40.3%).

Table 5: Dengue with warning signs by IgM Ab

P=0.0001 ;<0.05, S

Dengue with warning signs was more common in children with dengue IgM antibody test positive.

Table 6: Dengue with warning signs by NS1Ag category

P=0.004, <0.05, S

DW	Ns1 Ag		Total %	
	Positive (%)	Negative (%)		
Positive	15(60)	10 (40)	25	
Negative	9(24.3)	28(75.6)	37	
total	24(38.7)	38(61.3)	62	

Dengue with warning signs more common in children with NS1 Ag test positive.

Table 7: SEVERE DENGUE by IgM Ab category

positive	negative	
	0	
25 (98.4)	1(1.6)	26
5 (13.8)	31(86.2)	36
30(48.3)	32(51.7)	62
2 5 3	ositive 5 (98.4) (13.8) 0(48.3)	ositive negative 5 (98.4) 1(1.6) (13.8) 31(86.2) 0(48.3) 32(51.7)

P=0.001 ;< 0.05; S

Severe dengue is more common in children with dengue IgM antibody test positive.

Table 8: SEVERE DENGUE by NS1Ag

DSS	NS1Ag		Total (%)
	Positive (%)	Negative(%)	
positive	15 (57.6)	11 (42.3)	26 (100)
negative	9 (25)	27 (75)	36(100)
total	24 (38.7)	38 (61.3)	62(100)

P=0.009 ;< 0.05; NS

Severe Dengue more common in children with NSI Ag test positive.

Most common manifestations are pain abdomen, vomiting, arthralgias, body pains. More common bleeding manifestations in dengue with warning signs are skin bleeds and melena followed by epistaxis.

In severe dengue most common manifestations were skin bleeds (100%) (26/26) and melena (100%) (26/26).

DISCUSSION:

Dengue is an acute arboviral disease. It is probably one of the most important viral diseases in terms of human morbidity and mortality. The WHO says nearly 2.5 billion people , i.e., two fifths of the world population are now at risk from dengue and estimates that there may be 50 million cases of dengue infection worldwide every year³. The spectrum ranges from self limiting dengue fever to more severe forms of dengue fever with warning signs or severe dengue⁴. The problem of dengue has reached mammoth proportions in India since the first epidemic of clinical dengue like illness was recorded in Madras in 1780. It is compounded by the huge population, poor medical and diagnostic facilities and inadequate mosquito control³⁻⁶.

Among 62 cases studied 35 were males (56.45%) and 27. (43.55%) were females; male to female ratio is 1.3:1. In a study by Shubhankar Mishra etal $(2016)^7$, the male to female ratio was 3.4:1.Similar results were seen in a study by Pothapregada S et al $(2016)^8$, the male to female ratio was 1.2:1. In a study by Hemant Jain $(2016)^9$, out of 65 cases, males were 53% and females were 47%. In a study by Selvan T et al $(2017)^{10}$, the age and sex majority of the cases were in the age group of 10-18 years with 39.4% followed by 6-10 years with 34.5%.

In Shah G.S. et al $(2006)^{11}$, the mean age group was 8.3 yrs .In a study conducted by Ira shah et al, $(2005)^{12}$, the mean age group was 6.1yrs. In a study by Shubhankar Mishra et al $(2016)^7$, the mean age of patients was 8.7 yrs. In a study by Sahana KS et al $(2015)^{13}$, mean age of presentation was 8 y. In the study by Faridi et al $(2008)^{44}$, most of the cases (52.9%) were in the 10–12 year age group. In a study by Selvan T et al $(2017)^{10}$, male to female ratio was 1.7:1.In the present study also most of the reported cases were from the age group of 4-6 yr. In a study by Hemant Jain $(2016)^9$, maximum number of cases seen in the age group of 5-10 years (46%).

The present study the most common clinical presentation along with fever was pain abdomen (74%), vomiting (72.5%), arthralgia (4%), body pains (72.5%), poor intake (74%), hepatomegaly (74%).Similar observations were made in study conducted by Neeraja.M and Lakshmi.V. et al $(2006)^{14}$. In a study by Shubhankar Mishra et al $(2016)^7$, myalgia (76.8%) and abdominal pain (54.3%) were common symptoms. In a study by Sahana KS et al $(2015)^{13}$, vomiting (60.5%), pain abdomen (32%), headache (30.9%), myalgia (23.5%) and bleeding manifestations (16%) were the common presenting complaints.

In the present study tourniquet test positive in 32 patients out of 62 children constituting 51.6%. In the study by Faridi et al $(2008)^{15}$, positive tourniquet test was seen in 64.7%.

In present study common bleeding manifestation in severe dengue patients were skin bleeds i.e. petechiae (100%) and melena (100%) followed by epistaxis 73.8% and hematemesis 48.6%. Similar results were observed in a study by Shubhankar Mishra et al (2016)⁷, the most common bleeding manifestations in both severe and non-severe dengue were petechiae (22.1%). In study by Shah G.Set al (2006) ¹¹ common bleeding manifestations were skin bleeds (59%).In study by Gurdeep.S.Dhooria et al (2008)¹⁶, most common bleeding manifestations were petechiae in 85% followed by melena (6%), echymosis(2.5%) and epistaxis(2.5%).

In a study by Basuki PS et al(2010)using the WHO classification system 1997, 122 cases (84.1%) were classified as having non-severe dengue, of which 70 (48.3%) were classified as having dengue fever (DF), 39 (26.9%) as having dengue hemorrhagic fever (DHF) grade I, and 13 (9%) as having DHF grade II. Twenty-three (15.9%) were classified as having severe dengue, of which 16 (11%) were classified as having DHF grade III and 7 (4.8%) as having DHF grade IV.

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In a study by Sahana KS et al $(2015)^{13}$, number of cases classified as Dengue without warning signs , Dengue with warning signs (DW) and Severe Dengue (SD) were 48.1 %, 27.2 % and 24.7 % respectively. In a study by Hemant Jain $(2016)^{9}$ dengue without warning signs were 23%, dengue with warning signs were64%, and severe dengue were12% of cases.

In study by Gurdeep.S.Dhooria et al (2008)¹⁶92% of cases were dengue hemorrhagic fever, 7.4% cases presented in dengue shock syndrome.our study of 62 cases 8(13%) were probable dengue fever, 3 (4.8%) were dengue without warning signs, 25(40.3%) dengue with warning signs 26(40.9%) presented with severe dengue. Using the new dengue classification system 2009, In a study by Basuki PS et al(2010)¹⁷, 117 cases (80.7%) were classified as having non-severe dengue infection, of which 79 (54.5%) were classified as having dengue without warning signs and 38 (26.2%) were classified as having dengue with warning signs, while 28 (19.3%) were classified as having severe dengue infection. Based on the WHO TDR 2009 dengue guidelines, in a study by Shubhankar Mishra, et al (2016)⁷, the total number of cases analyzed was 97, out of which84 (86.59%) were categorized as cases of non-severe dengue which included both undifferentiated fever and dengue fever (DF) (both with and without warning signs) and 13 (13.40%) were cases of severe dengue (DHF grades 1–4).

In this study Ns1Ag test was positive 29.2% cases. Similar observation seen in study by B.Mustafa MPH et al $(31.2\%)(2010)^{18}$. In our study there is strong correlation present between NS1Ag positivity and severe dengue and dengue with warning signs. In a study by Shubhankar Mishra et al. $(2016)^7$, the majority of the patients were positive for NS1 followed by IgM. In a study by Pothapregada S et al $(2016)^8$, Dengue IgG antibody was positive in 17 cases (6.5%) and among them, severe dengue infection was seen in 16 cases (94.1%). In study by Hemant Jain $(2016)^9$, NS 1 antigen was positive in 80% of cases; IgM and IgG was positive in 52% and 36% cases respectively.

CONCLUSION:

The detection of IgM dengue antibodies by capture ELISA & NS1 Ag were helpful for diagnosis of acute dengue virus infection. The serological diagnosis of dengue fever has a role in categorizing primary and secondary infection and it also serves as a predictor of disease progression and mortality especially in severe dengue.

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