A retrospective study on the epidemiological and clinical characteristics of SARS-CoV-2 infection among the affected pediatric population attending a tertiary care hospital in **Eastern India**

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the all age groups, including the pediatric population. Viral respiratory infections in children

may act as a source of infection especially those with asymptomatic SARS-CoV-2 infections.

Aims and Objectives: The aim is to analyze the SARS-CoV-2 characteristics in pediatric age

group. Materials and Methods: Ninety seven children neonate to 5 years age tested positive

by RT-PCR were included in the study. Detailed data on socio-demography, epidemiology,

clinical characteristics, laboratory findings, comorbidities, management, and outcome of the study population were collected. Results: Sex distribution showed 47 (48.4%) male and 50 (51.5%) female. Among these 97, 5 children (5.1%) were <28 days of age, 30

children were (31%) neonate to infant age, and 62 children were (63.9%) above 1 year

but <5 years of age. The group was divided into two---one with 56 children (57.7%) who were discharged, 2nd with 41 (42.3%) hospitalized children. 1st group was asymptomatic (64.3%)/mildly symptomatic (35.7%). In 2nd group 36.5% were moderately symptomatic, 29.2% were with severe complications, and 34.3% of children were critical--22 (52.6%) required PICU admission--17 children (41.4%) required mechanical ventilation and 36 (87.8%) required respiratory support. Conclusion: Reports state that COVID-19 infection is mild in

the pediatric population but our findings suggest that some children can develop severe/

critical infection requiring hospitalization, respiratory support, and mechanical ventilation.

Key words: COVID-19; Disease severity; Inflammation; Pediatric population;

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ABSTRACT

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Background: Novel coronavirus infections cause major morbidity and mortality globally in

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INTRODUCTION

Retrospective study

COVID-19 infection is caused by the severe acute respiratory syndrome coronavirus-2 (SARS-CoV-2) virus, a beta virus family Coronaviridae¹, has high lethality and transmissibility affecting people all age groups.² Surprisingly, children are affected in very small proportion (1–10% positive cases worldwide).³ Reviews have described

mild to moderate illness in children with full recovery in 1–3 weeks.⁴ However, children may not show any signs of infection, delaying its diagnosis, leading to complications and community spread.⁵ As the risk of intrauterine transmission is low, close contact with a source is the probably the cause of positive results in neonates.^{6,7} This retrospective study has been designed to lay emphasis on the understanding of different socio-demographic

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and clinical risk factors associated with COVID positive children in and around Burdwan from May 2020 to December 2020.

Aims and objectives

The aim is to analyze the different socio-demographic and clinical risk factors affecting SARS-COV-2 infection in the pediatric age group.

MATERIALS AND METHODS

Patient selection

A retrospective study was conducted in a tertiary care COVID referral hospital at Burdwan town of West Bengal with all children and newborns attending its outpatients department for COVID testing for 8 months--May 2020 to December 2020. A total of 97 children from age <1 month to 5 years who were tested COVID-19 positive by RT-PCR at Burdwan Medical College during the span of this period were included in the study.

Sample collection

Nasopharyngeal and oropharyngeal swabs were collected into VTMs and transported immediately to the COVID-19 referral laboratory of this institution for RT-PCR test.

Data collection

Detailed data on socio-demography, epidemiology, clinical characteristics, laboratory findings, co-morbidities, management, and outcome of our study population were collected by communicating with the guardians of the patients and also from the available medical records. All data were recorded in a pre-piloted health counseling form. Data of age groups and disease severity were extracted wherever available either from the form or from discussion with the guardians. Disease severity was classified as per the national guidelines and as discussed in Table 1. Symptomatic analysis of the patients were done and recorded in the health counseling form.

Data extraction

We recorded whether fever, cough, cold, sore throat, respiratory distress, or skin manifestations were present. For diagnosis of hyper inflammatory Syndrome/PIMS-TS we followed the diagnostic criteria as published by the World Health Organization. In patients with severe disease requiring admission in the hospital, the following data were collected-admission in pediatric intensive care units (PICUs), requirement of mechanical ventilation, presence of hypoxia (oxygen saturation <92%) and diagnosis of multi-organ failure.

Ethical considerations

For the study, permission from the Institutional Ethics Committee (IEC) of Burdwan Medical College and Hospital was taken (BMC/IEC/133, dated March 15, 2021). Our study included reports of all clinical examination, medical questionnaires, personal, and family history of the patients as provided by the patients/guardian of the patients.

Data analysis

Statistical analysis by Fisher exact test, or Mantel-Haenszel correction was done to determine if there was any association between different variables. Odds ratios with confidence intervals of 95% were calculated. Data were analyzed with SPSS v16.0 (Table 1).

RESULTS

A total of 97 COVID positive children who attended the outdoor unit of Burdwan Medical College and Hospital during May 2020 to December 2020 were selected for this study. Distribution by sex showed that 47 (48.4%) were male and 50 (51.5%) were female (OR 1.47, 95% CI 1.02–3.18, P=0.47). Among these 97, 5 (5.1%) were <28 days of age,

Table 1: Crit	teria for predicting disease severity
Asymptomatic	 Signs or symptoms absent. Chest X-ray normal
Mild	 No other suggestive condition present. Symptoms of upper respiratory tract infectioncough, sore throat, nasal congestion, malaise and headache with fever but no breathlessness (normal saturation).
Moderate	 Chest X-ray showing no signs of pneumonia Dyspnea, fever, cough, and SpO₂<94%;
	respiratory rate<24/min. 2. Increased respiratory rate in<2 months>60/min;
Severe	2-11 months≥50/min; 1–5 years≥40/min. 1. Oxygen saturation SpO ₂ <90%
	2. Signs of severe respiratory distress (apnea gasping, and head nodding)
	3. Child with severe cough/respiratory distress with at least one of the followingcentral
	cyanosis, grunting, and chest in-drawing.4. Signs of pneumonia with any of the followingextreme lassitude, lethargy,
	convulsions, or unconsciousness. 5. Signs of pneumoniachest in-drawing,
	increased respiratory rate<2 months≥60; 2–11 months≥50; 1–5 years≥40.
Critical	 6. Need for any respiratory support 1. Patient in intensive care unit with acute
	respiratory distress syndrome. 2. Altered mental state; bradycardia or
	2. Altered mental state, bradycardia of tachycardia (HR 160 bpm in infants and HR 150 bpm in children); prolonged capillary refill (>2 s) or weak pulse; tachypnea; mottled or cool skin; petechial or purpuric rash; high lactate level; reduced urine output; hyperthermia; or hypothermia
	 Multi-organ failure Shock, encephalopathy, myocardial injury
	coagulation dysfunction, features of acute renal failure.

30 (31%) were between neonate to infancy, and 62 (63.9%) were above 1 year to <5 years of age. The total group was divided into two sub-groups, first group with 56 children (57.7%) who were discharged at home while the second group with 41 (42.3%) children who were hospitalized. About 53.6% children belonging to the discharged group were male whereas in case of the hospitalized group the male prevalence was less, 41.4%. About 91% and 90.2% of the children were of Bengali ethnic origin among the discharged and hospitalized groups, respectively. Among the discharged group, 33 (58.9%) children had at least one SARS-CoV-2 infected family member, whereas among the hospitalized group, 25 (60.9%) children had an infected family member. The mean+SD for the Ct values (in real time PCR for SARS-CoV-2 RNA) of children belonging to the discharged and hospitalized groups were 25.6+2.7 and 23.8+3.2, respectively. 12 (21.5%) children belonging to the discharged group and 13 (31.7%) children belonging to the hospitalized group had co-morbidities such as cardiac, renal, respiratory, and neuro-muscular diseases. 22 (52.6%) of the hospitalized children required PICU admission, 17 (41.4%) of them required mechanical ventilation, and 36 (87.8%) hospitalized children required respiratory support. CT scan infective changes including ground glass opacity and focal consolidation were observed in 10.7% of the discharged patients and in 65.8% of the hospitalized patients. Most children belonging to the discharged group were asymptomatic (64.3%) and the rest 35.7% were mildly symptomatic. Among the hospitalized children 36.5% were moderately symptomatic, 29.2% presented severe complications and 34.3% had a critical presentation. The mean CRP value was significantly higher in case of the hospitalized children compared to those who were discharged. The mean D-Dimer and ESR values were more or less similar for both the groups. Leukopenia was observed in case of 19.6% of the discharged children and 56.1% of the hospitalized children. Lymphopenia was observed in 12.5% of the discharged children and 29.2% among those hospitalized. Only one child out of the 56 children discharged and two children out of those 41 children hospitalized died during the course of this study. The detailed analysis is presented in Table 2.

Among the discharged group with 56 children, fever was recorded in 16 children (28.5%), sore throat among 12 (21.4%), body ache among 10 (17.8%), dyspnea among 9 (16.1%), headache among 4 (7.1%), vomiting among 8 (14.3%), diarrhea among 5 (8.9%), dry cough among 17 (30.3%), thoracic pain among 3 (5.35%), febrile convulsions among 6 (10.7%), respiratory distress among 8 (14.3%), otitis among 3 (5.35%), muscle pain among 11 (19.6%), joint pain among 7 (12.5%), and rhinorrhea among 12 (21.4%). Two children were hypoxemic at presentation. Among children with vomiting, one had blood mixed vomiting.

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Among the hospitalized group with 41 children, 23 (56.1%) children reported fever and 15 (36.5%) reported sore throat. Among the other symptoms reported body ache was observed among 8 (19.5%), dyspnea among 12 (29.2%), headache among 6 (14.6%), vomiting among 10 (17.1%), diarrhea among 6 (14.6%), dry cough among 12 (29.2%), thoracic pain among 5 (12.2%), febrile convulsions among 6 (14.6%), respiratory distress among 26 (54.6%), otitis among 2 (4.9%), muscle pain among 15 (36.5%), joint pain among 10 (24.4%), and rhinorrhea among 22 (53.6%).

The data related to these clinical symptoms and manifestations are graphically represented in Figure 1.

DISCUSSION

This article adds to the previous knowledge of clinical characteristics and outcomes of SARS-CoV-2 infection in pediatric age group. A small population of 97 COVID infected children in the Burdwan District of West Bengal, India. This study, therefore diagnoses the prevalence of positive cases in children with characteristics based on socio-demographic variables, co-morbidities, disease severity, clinical manifestations, laboratory parameters, CT-Scan, and oxygen support. The patients in our study showed higher rate of PICU admission per hospitalization (22.6%), which may be due to a number of social factors negatively influencing the health status. Various reports suggest that the infection causes a milder impact on children, but our findings suggest that some children develop severe/critical disease and require immediate hospitalization and in some cases PICU admission, respiratory support, and mechanical ventilation. It has been found that these children show higher level of the inflammatory marker like CRP which probably lead to higher rate of ARDS. (Serum levels of different inflammatory cytokines linked to ARDS were,

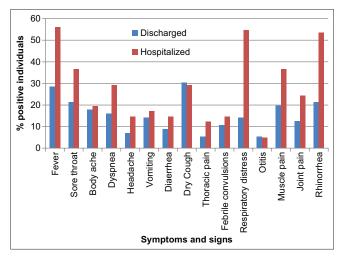


Figure 1: Schematic representation of different symptoms associated with the COVID positive children

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Characteristics	Discharged at home (n=56)	Hospitalized (n=41)	P value
Age			
<1 month	4 (7.3%)	1 (2.4%)	0.19
1 month to 1 year	8 (14.2%)	22 (53.6%)	
1 year to 5 years	44 (78.5%)	18 (44%)	
Sex			
Male	30 (53.6%)	17 (41.4%)	0.73
Female	26 (46.4%)	24 (58.6%)	
Ethnicity			
Bengali	51 (91%)	37 (90.2%)	0.02
Non-Bengali	5 (9%)	4 (9.8%)	
Family members infected	(),		
Yes	33 (58.9%)	25 (60.9%)	0.21
Νο	23 (41.1%)	16 (39.1%)	
Ct value for COVID RT-PCR	Mean=25.6+2.7	23.8+3.2	0.32
Disease severity			
Asymptomatic	36 (64.3%)	0	0.003
Mild	20 (35.7%)	Ő	0.000
Moderate	0	15 (36.5%)	
Severe	0	12 (29.2%)	
Critical	0	14 (34.3%)	
D-Dimer level (ug/ml)	Mean=0.9+0.26	0.83+0.34	0.41
CRP mg/dl	Mean=1.8+1.5	6.3+2.6	0.41
ESR mm/h	Mean=26+2.5	28+1.3	0.02
	Wear1-20+2.5	20+1.3	0.22
Leukopenia	44 (40,00())	00 (50 40())	0.00
Yes	11 (19.6%)	23 (56.1%)	0.06
No	45 (80.4%)	18 (43.9%)	
Lymphopenia	- ////		
Yes	7 (12.5%)	12 (29.2%)	0.12
No	49 (87.5%)	29 (70.8%)	
CT scan-infective changes			
Yes	6 (10.7%)	27 (65.8%)	0.033
No	50 (89.3%)	14 (34.2%)	
Respiratory support			
Yes	0	36 (87.8%)	0.025
No	56 (100%)	5 (12.2%)	
Mechanical ventilation			
Yes	0	17 (41.4%)	0.41
No	56 (100%)	24 (58.6%)	
Co-morbidities			
Yes	12 (21.5%)	13 (31.7%)	0.81
No	44 (78.5%)	28 (68.3%)	
Pediatric intensive care unit admission	(/	- ()	
Yes	0	22 (53.6%)	0.04
No	56 (100%)	19 (46.4%)	0.04
Outcome			
Cured	53 (94.6%)	39 (95.1%)	0.43
Dead	1 (1.8%)	2 (4.9%)	0.40

Table 2: Socio-demographic characteristics and disease severity of the COVID positive children at

however, not estimated.) Different articles suggest that children co-morbidities have critical COVID-19 infection; but no such direct correlation could be found in our study. About 33.25% of our patients gave the history of coming in close proximity with an infected person. This high rate of community spread can be due to high population density in the region of our study, illiteracy and also failure to observe the requisite precautions of the disease.

A similar retrospective study⁸ was done in Morocco on 74 COVID affected children. They found that females were affected more by the infection than males like other

studies.9 But in our study, no such sex predilection was noted (47 affected children were male and 50 were female - hospitalized and discharged taken together).

We found in our study that symptoms related to the respiratory and digestive systems were predominantly reported, similar to other studies.⁷ In our study, we found fever and cough was the commonest symptom in children. Different studies have also reported so^{10,11} (40-60% in children). In our study, we also found that diarrhea and vomiting was very commonly complained by the children which is similar to other studies.¹²

We also found leukopenia and not leukocytosis in 19.6% of the discharged children and 56.1% of the hospitalized children which has also been reported in other studies.¹³ Lymphopenia was observed in 12.5% of the discharged children and 29.2% among those hospitalized, similar to a study done in Morocco on children who reported an incidence of 3% in their study population.^{8,14}

A Chinese study¹⁵ had reported that infection in nearly 90% children is either asymptomatic or mild to moderate — severe infection being very less. Moroccan study also substantiated this finding. Even in our study, most children belonging to the discharged group were asymptomatic (64.3%) and the rest 35.7% were mildly symptomatic. Among the hospitalized children 36.5% were moderately symptomatic, 29.2% presented severe complications, and 34.3% had a critical presentation.

Similarly, mortality is very much less in the children compared to adults. Only one child out of the 56 children discharged and two children out of those 41 children hospitalized died during the course of this study. This fact is also reported in different studies^{16,17}

Hence, this study with a small population of children (97) during the time span of May 2020 to December 2020 developed mild COVID infection. This is in concordance with other studies done worldwide in this time span. The children were all discharged healthy and only a small percentage developed complication and only three died. Thus, based on our findings, it can be inferred that COVID 19 is a mild infection in children with very few life-threatening complications.

Limitations of the study

Sample size was not adequate as this retrospective study was conducted on a rapid emerging novel corona virus infection for a short period of time in a rural tertiary teaching hospital.

CONCLUSION

Here, we have presented a small retrospective observational study with a limited sample size attending a single, medical center, leading to type II statistical error, a major limitation to our study. However, as COVID-19 is a relatively new disease with limited number of data resources available from the infected pediatric population, our study will surely provide a better understanding of the infection.

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Authors Contribution:

TB- Concept and design of the study, prepared first draft of manuscript; NC- Reviewed the literature and manuscript preparation; RM- Statistical analysis and interpretation; AC-laboratory test and interpretation; SC- Concept and design of the study, prepared first draft of manuscript, preparation of manuscript, and revision of manuscript

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