

BRIEF REPORT

Initial Observations of COVID-19 in US Children

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Coronavirus disease (COVID-19) has affected children differently from adults worldwide. Data on the clinical presentation of the infection in children are limited. We present a detailed account of pediatric inpatients infected with severe acute respiratory syndrome coronavirus 2 virus at our institution during widespread local transmission, aiming to understand disease presentation and outcomes. A retrospective chart review was performed of children, ages 0 to 18 years, with a positive polymerase chain reaction test for severe acute respiratory syndrome coronavirus 2 on nasopharyngeal specimens admitted to our hospital over a 4-week period. We present clinical data from 22 patients and highlight the variability of the presentation. In our study, most children presented without respiratory illness or symptoms suggestive of COVID-19; many were identified only because of universal testing. Because children may have variable signs and symptoms of COVID-19 infection, targeted testing may miss some cases.

ABSTRACT

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Children account for <5% of the cases of severe acute respiratory syndrome coronavirus 2 infections in the United States to date.^{1,2} Although some general epidemiological data of coronavirus disease (COVID-19) in children have been published,³⁻⁶ there is limited reporting of the actual clinical presentation. We present a detailed account of pediatric patients who presented to our institution during the early stages of the COVID-19 pandemic to provide a better understanding of the disease presentation and outcomes in children.

METHODS

A retrospective review was performed of patients, ages 0 to 18 years old, admitted to our inpatient pediatric service at a children's hospital in Brooklyn, New York from March 18 to April 15, 2020, who tested positive by polymerase chain reaction (PCR) for SARS-CoV-2 virus on a nasopharyngeal specimen. Patients were identified using the hospital's daily log that listed all institutional testing results for SARS-CoV-2. Our initial testing strategy was in accordance with Centers for Disease Control and Prevention guidelines, recommending testing if there were symptoms of fever, cough, and shortness of breath, travel to high risk countries, or close contact with a confirmed case. As the incidence of infection increased, in the latter half of our study, from March 27 onwards, we implemented PCR testing for all admitted patients irrespective of symptomatology. This study was approved by the institutional internal review board for expedited review.

RESULTS

Of the 22 patients who tested positive, 55% were male (Table 1). Children ranged in age from 11 days to 18 years. Infants <1 year of age made up 45% of hospitalizations. No patient had a travel history, and 6 of 22 (27%) had confirmed SARS-CoV-2 exposure. Six patients had underlying comorbidities (3 with malignancy, 2 with chronic lung disease, and 1 with cardiac disease). The majority of patients, 18 of 22 (82%), were admitted to the hospital within 3 days of

Table 1 Characteristics of Hospitalized Pediatric Patients With COVID-19

Characteristic (N = 22)	No.	%
Sex		
Male	12	55
Female	10	45
Age, y		
Distribution		
<1	10	45
1-6	4	18
7-12	3	14
13-18	5	23
Presenting symptoms		
Fever	15	68
Any respiratory symptom	9	41
Difficulty breathing	6	27
Nasal congestion	5	23
Cough	4	18
Fatigue	6	27
Seizures	2	9
Headache	1	4
Duration of symptoms before admission, d		
Asymptomatic	2	9
<1	3	14
1-3	13	59
>3	4	18
Known COVID-19 contact		
At home	3	14
Outside of home	3	14
None known	16	72
Underlying medical conditions	6	27
Other admitting diagnosis and COVID-19 positivity	7	32
Respiratory support		
Noninvasive ventilation	3	14
Mechanical ventilation	4	18
None	15	68
Viral coinfection/total tested	2/7	29
Chest radiograph abnormalities/total imaged	5/11	45
Laboratory abnormalities/total tested		
CRP >1 mg/L	8/10	80 ^a
Procalcitonin > 0.5 ng/mL	6/7	86 ^b
Absolute lymphocytes <1500/ μ L	7/22	32
Transaminitis	2/7	29

Underlying medical conditions were as follows: malignancy, 3; bronchiectasis, 1; cardiac (patent ductus arteriosus-closed ventricular septal defect), 1; prematurity or chronic lung disease, 1. Other admitting diagnoses were 1 each of perforated appendix, urinary tract infection, cellulitis, septic arthritis, cardiac arrest, purulent otorrhea, and myositis. Patients also tested positive for SARS-CoV-2. Viral coinfection indicates positive for enterovirus/rhinovirus. Respiratory support (noninvasive ventilation) includes nasal cannula, high flow oxygen, and bilevel positive airway pressure.

^a 4 of 8 with codiagnoses.

^b 3 of 7 with underlying medical condition.

symptom onset. No patient died during the study period.

The most common clinical presentation was fever without a source in otherwise healthy infants (5 of 22; 23%), with age range 11 to 35 days. All 5 patients had a sepsis evaluation, including cerebrospinal fluid analysis, received empirical antibiotics, and were discharged from the hospital once the bacterial cultures were negative within 48 to 72 hours.

Only 9 (41%) patients presented with a respiratory illness, and 7 (32%) required respiratory support. Four patients needed mechanical ventilation; 2 of these patients had underlying pulmonary disease, a teenager with bronchiectasis and a 1-year-old with chronic lung disease due to prematurity. Both progressed within 6 to 72 hours from high flow oxygen support to ICU admission and intubation. Of the two other patients who required intubation, 1 had cerebral palsy and status epilepticus and the second child was otherwise healthy and presented in cardiac arrest.

Most patients with respiratory illness were managed with supportive therapy and antibiotics as indicated. However, three patients admitted to the PICU and on mechanical ventilation qualified for compassionate use of remdesivir. The drug was only available for patients with documented infection and respiratory deterioration requiring mechanical ventilation without concomitant liver or kidney disease. All three of the patients treated with remdesivir were eventually extubated.

Two patients had neurologic abnormalities: an 11-year-old healthy boy presented with fever, headache, confusion, and seizure. His cerebrospinal fluid showed mild pleocytosis (white blood cell count: 16, red blood cell count: 921), protein 92 mg/dL, glucose 97 mg/dL, the cerebrospinal fluid PCR panel was negative, and he had an abnormal EEG (diffuse cerebral dysfunction); he improved, without short-term sequelae, within 48 hours. A second patient, a 12-year-old girl with cerebral palsy, developed new onset seizures after several days of fever and cough, requiring mechanical ventilation. She

improved to baseline after 18 days in the hospital.

Three patients with malignancies were hospitalized. One presented with mild sore throat and fever; the second was asymptomatic and admitted for routine chemotherapy. The third patient, a teenager, had bilateral pneumonitis and hypoxia and required oxygen therapy for 3 days.

In terms of laboratory abnormalities, lymphopenia was noted in 32%, and an elevated procalcitonin or C-reactive protein were present in the majority of patients in whom the tests were performed. Abnormal chest radiograph findings, with bilateral opacities, were noted in 5 of 11 patients (Table 1). Viral coinfection was detected in 2 of 7 tested for other viruses.

During the second half of the study period, a positive PCR result was noted in 7 patients (32%) who were hospitalized for non-COVID-19-related symptoms. Four patients had documented bacterial infections, and one was diagnosed with appendicitis (Table 1). The other two presented with illnesses of unclear etiology. One had inflammation of the forearm muscles with no abscess formation but fever and elevated inflammatory markers, was treated with antibiotics, and did well. A third patient, a 6-month-old boy, presented after cardiac arrest at home with no known underlying diseases; his echocardiogram showed severely depressed ventricular function, and his chest radiograph at the time of admission was normal.

Two patients were completely asymptomatic at the time of admission but were positive by PCR; one was admitted for social reasons and the other for routine chemotherapy as mentioned above.

DISCUSSION

In our study, hospitalized pediatric patients with COVID-19 had a wide spectrum of presentation, and few displayed the classic respiratory symptoms associated with this disease in the adult population; only 41% of admitted children had respiratory tract illness. These findings differ from the description of the disease in several initial

studies out of China, where the major presentation was a respiratory illness of varying severity,⁴⁻⁶ but are similar to findings from a more recent meta-analysis.⁷ Almost half of our cohort was aged <1 year, and half of those were <6 weeks of age presenting with fever alone, necessitating an evaluation for sepsis. Our findings again reveal differences between the reports from China, where in a large study of 171 children, only 18% were aged <1 year, and the median age of presentation was 6.7 years.⁵

Our initial testing strategy was according to the federal and local guidelines that recommended PCR testing for the symptoms of fever, cough, and shortness of breath or travel to certain countries or close contact with a confirmed case. With the implementation of our universal screening strategy of all admitted pediatric patients, we identified 9 (41%) patients with COVID-19 who would have been missed because they did not meet the then-recommended criteria for testing. For the patients admitted with alternate diagnoses, it is not clear if and how significant a role SARS-CoV-2 had in their illness. Two patients presented with encephalitis with no alternative etiology; it is possible that SARS-CoV-2 was the cause in both cases.

Our strategy also led to documenting asymptomatic infection in two patients, one of whom was immunocompromised and needed chemotherapy to be postponed based on the test result. Finally, the youngest infant in our cohort, an 11-day-old, was born to a mother who was well and family members were asymptomatic, suggestive of asymptomatic transmission in the home. Only a minority of our patients (28%) had documented confirmed viral exposure, highlighting that the infection rates at a given time in a particular city should drive the strategies of pediatric testing, rather than confirmed contact alone. This finding also is unexpected because several of the reports from China³⁻⁵ describe the vast majority of transmission in children from family clusters.

CONCLUSIONS

Early experience at our hospital shows that most hospitalized pediatric patients did not

present with the classic symptoms attributed to COVID-19 and the majority did not have household exposure to the infection, thereby presenting atypically from what is seen in adults and the reported pediatric experience from China. Guidelines to test pediatric patients need to be broadened and take into account that patients presenting with other illnesses may also be positive for COVID-19. Testing of all hospitalized patients will not only identify cases early in the course of their admission process but will also help prevent inadvertent exposure of other patients and health care workers, assist in cohorting infected patients, and aid in conservation of personal protective equipment.

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