#### **ORIGINAL ARTICLE**



# Worse Hospital Outcomes for Children and Adults with COVID-19 and Congenital Heart Disease

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#### Abstract

The aim of the current study is to investigate hospitalization outcomes of COVID-19 positive children and adults with moderate or severe congenital heart disease to children and adults without congenital heart disease. Retrospective review using the Vizient Clinical Data Base for admissions of patients with an ICD-10 code for COVID-19 from April 2020 to March 2021. Admissions with COVID-19 and with and without moderate or severe congenital heart disease (CHD) were stratified into pediatric (<18 years) and adult ( $\geq$ 18 years) and hospital outcomes were compared. There were 9478 pediatric COVID-19 admissions, 160 (1.7%) with CHD, and 658,230 adult COVID-19 admissions, 389 (0.06%) with CHD. Pediatric admissions with COVID-19 and CHD were younger (1 vs 11 years), had longer length of stay (22 vs 6 days), higher complication rates (6.9 vs 1.1%), higher mortality rates (3.8, 0.8%), and higher costs (\$54,619 vs 10,731; p < 0.001 for all). Adult admissions with COVID-19 and CHD were younger (53 vs 64 years, p < 0.001), had longer length of stay (12 vs 9 days, p < 0.001), higher complication rates (8 vs 4.8%, p = 0.003), and higher costs (\$23,551 vs 13,311, p < 0.001). This appears to be the first study to report the increased hospital morbidities and costs for patients with CHD affected by COVID-19. Our hope is that these findings will help counsel patients moving forward during the pandemic.

**Keywords** COVID-19  $\cdot$  Hospital outcomes  $\cdot$  Congenital heart disease  $\cdot$  Fontan  $\cdot$  Tetralogy of Fallot  $\cdot$  Transposition of the great arteries

## Introduction

As of July 2021, there were 183,696,230 cases of coronavirus disease 2019 (COVID-19) globally, with 3,975,227 deaths [1]. Patients who have underlying cardiovascular conditions such as hypertension and coronary heart disease have additional morbidity and mortality for COVID-19 [2, 3]. Patients with moderate and severe congenital heart disease (CHD) have empirically been considered higher risk from COVID-19 [3, 4] though, due to their relatively small

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population, there are sparse data regarding the outcomes of hospitalized patients with moderate and severe CHD along with COVID-19. The purpose of this study was to use a national discharge database to analyze and report hospital outcomes for COVID-19 in the setting of moderate and severe CHD compared to those without CHD.

### Methods

The Vizient Clinical Data Base is an analytic platform for performance improvement populated by hundreds of health systems and community hospitals in the USA, including nearly all academic medical centers. The database includes demographics, mortality, length of stay (LOS), complications, readmission rates, resource utilization, and other information. After approval from the University of Arizona Institutional Review Board, which waived the need for informed consent for this retrospective review of deidentified data, we queried the database

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for hospital discharge data from April 2020 to March 2021 for all admissions with an ICD-10 code for moderate or severe CHD, as defined by the 2018 Adult CHD guidelines (Table 1) [5], and the diagnostic code for COVID-19 (U07.1), which went into effect April 1, 2020. The Adult CHD guidelines determined CHD severity by native anatomy, surgical repair, and current physiology [5]. Diagnoses of an isolated atrial septal defect, congenital heart block, isolated levocardia, tortuous aortic arch, anomalous subclavian artery, and any "NOS" cardiac diagnoses were excluded. The comparison group (No CHD) consisted of admissions with no ICD-10 codes for CHD and with the diagnostic code for COVID-19. The groups were further divided into pediatric (<18 years) and adult ( $\geq$ 18 years). Comparison of outcomes between acyanotic and cyanotic lesions was also performed. Demographics, LOS, rate of the presence of complications listed in the Vizient Clinical Data Base [6], in-hospital mortality, and total hospital costs were collected. To determine if there was a difference in admission severity of illness or mortality risk between the groups, queries were preformed from Vizientdefined "Acuity Scale Mortality" (severity of illness at admission) and "Relative Expected Mortality" (mortality risk at admission). These were converted to dichotomous values of Low/normal (comparable or less risk) and High (above normal risk) for comparison. Continuous, normally distributed data are presented as mean ± standard deviation, non-normally distributed as median (interquartile range), and categorical data as number (%). Comparisons were made between groups using the *t*-test for normally distributed data, the Kruskal-Wallis test for non-normally distributed data, and the  $\chi^2$  test for categorical data. A p-value < 0.05 was considered statistically significant.

Table 1 ICD-10 codes for moderate and severe congenital heart disease, as defined by the 2018 Adult Congenital Heart Disease guidelines

Moderate congenital heart disease	ICD-10 codes
Partial or total anomalous pulmonary venous return	Q26.2, Q26.3, Q26.4
Anomalous left coronary artery from the pulmonary artery	Q24.5
Atrioventricular septal defect	Q21.2
Aortic stenosis (congenital)	Q23.0
Mitral stenosis (congenital)	Q23.2
Coarctation of the aorta	Q25.1, Q25.42
Cor triatriatum	Q24.2
Ebstein anomaly	Q22.5
Subpulmonary stenosis	Q24.3
Patent ductus arteriosus	Q21.4, Q25.0
Pulmonary regurgitation	Q22.2
Pulmonary stenosis	Q22.1
Peripheral pulmonary stenosis	Q25.6, Q25.7, Q25.71
Sinus of Valsalva fistula	Q25.4
Subaortic stenosis	Q24.4
Supravalvar aortic stenosis	Q25.3
Tetralogy of Fallot	Q21.3
Double aortic arch	Q25.45
Complex congenital heart disease	ICD-10 codes
Double outlet right ventricle	Q20.1
Interrupted aortic arch	Q25.21
Single ventricle (DILV, TA, HLHS, etc.)	Q20.4, Q22.4, Q22.6, Q23.4, Q25.2, Q25.29, Q25.41
Pulmonary atresia	Q22.0, Q25.5
Transposition of the great arteries (d-, l- or congenitally corrected)	Q20.3, Q20.5
Truncus arteriosus	Q20.0
Criss cross heart, heterotaxy	Q20.6
Eisenmenger	127.83
Double outlet left ventricle	Q20.2

DILV double inlet left ventricle; HLHS hypoplastic left heart syndrome; TA tricuspid atresia

Statistical analyses were performed using SPSS 27 (IBM Corporation, Armonk, New York, USA).

## Results

There were 9478 total pediatric COVID-19 admissions, 160 (1.7%) with CHD, and 658,230 total adult COVID-19 admissions, 389 (0.06%) with CHD. Demographics and hospital outcomes are shown in Table 2. Pediatric admissions with COVID-19 and CHD were younger, had longer LOS, higher presence of complications listed in the Vizient Clinical Data Base, higher mortality rates, and higher costs (Table 2). There were no sex-based or race/ethnicity-based differences. Adult admissions with COVID-19 and CHD were younger, had longer LOS, higher presence of complications listed in the Vizient Clinical Data Base, and higher costs (Table 2). There were no sex-based or race/ethnicity-based differences listed in the Vizient Clinical Data Base, and higher costs (Table 2). There were no differences in in-hospital mortality rates or sex-based

or race/ethnicity-based differences. The specific CHD diagnoses for the Pediatric and Adult groups are listed in Table 3. There were no differences in acute hospital outcomes (LOS, presence of complications listed in the Vizient Data Base or mortality) between cyanotic and acyanotic diagnoses (Table 4). Pediatric admissions with CHD had higher severity of illness and mortality risk at admission, but adults with CHD had lower severity of illness at admission with no difference in mortality risk (Table 5).

#### Discussion

COVID-19 is a rapidly evolving global pandemic. While non-congenital cardiovascular comorbidities have been identified as risk factors for poor outcomes [2], to the best of our knowledge, ours is the first national study to demonstrate worse hospital outcomes for children and adults

Table 2 Demographics and hospital outcomes for pediatric (<18 years old) and adult ( $\geq$ 18 years) admissions for COVID-19 infection with and without moderate or severe congenital heart disease (CHD)

Pediatric (<18 years)	CHD (n = 160)	No CHD (n=9,318)	р
Female $(n, \%)$	77 (48)	4,770 (51)	0.442
Race/ethnicity (n, %)			
White	72 (45)	4,146 (45)	0.899
Black	40 (25)	2,054 (22)	
Hispanic	53 (33)	2,983 (32)	
Asian	3 (2)	223 (2)	
LOS (d)	$22.2 \pm 42.7$	$6.3 \pm 20.6$	< 0.001
Complications $(n, \%)$	11 (6.9)	101 (1.1)	< 0.001
Death ( <i>n</i> , %)	6 (3.8)	79 (0.8)	< 0.001
Direct costs (\$)	$54,619 \pm 124,413$	$10,731 \pm 39,952$	< 0.001
Adult (≥18 years)	CHD	No CHD	р
	( <i>n</i> =389)	(n = 657, 841)	
Age (y)	53 (35, 65)	64 (50, 76)	< 0.001
Female (n, %)	192 (49)	312,510 (48)	0.465
Race/ethnicity (n, %)			
White	236 (61)	364,706 (55)	0.121
Black	70 (18)	143,500 (22)	
Hispanic	69 (18)	127,274 (19)	
Asian	17 (4)	23,057 (4)	
LOS (d)	$11.6 \pm 14.5$	$8.7 \pm 11.0$	< 0.001
Complications $(n, \%)$	31 (8)	31,385 (4.8)	0.003
Death ( <i>n</i> , %)	41 (10.5)	79,594 (12.1)	0.346
Direct costs (\$)	$23,551 \pm 44,503$	$13,311 \pm 25,891$	< 0.001

Data are presented as n (%), mean  $\pm$  standard deviation or median (interquartile range)

Data from the Vizient Clinical Data Base used by permission of Vizient. All rights reserved

CHD congenital heart disease; LOS length of stay; No CHD no congenital heart disease

 Table 3
 Congenital heart defect

 diagnoses for the pediatric and
 adult groups

	Pediatric CHD	Adult CHD	
	n	n	
Moderate congenital heart disease			
Partial or total anomalous pulmonary venous return	6	15	
Anomalous left coronary artery from the pulmonary artery	9	89	
Atrioventricular septal defect	11	14	
Aortic stenosis (congenital)	1	14	
Mitral stenosis (congenital)	1	6	
Coarctation of the aorta	16	26	
Cor triatriatum	0	5	
Ebstein anomaly	6	16	
Subpulmonary stenosis	1	2	
Patent ductus arteriosus	61	34	
Pulmonary regurgitation	1	5	
Pulmonary stenosis	4	14	
Peripheral pulmonary stenosis	27	34	
Sinus of Valsalva fistula	0	0	
Subaortic stenosis	2	13	
Supravalvar aortic stenosis	0	2	
Tetralogy of Fallot	22	31	
Double aortic arch	1	4	
Complex congenital heart disease			
Double outlet right ventricle	12	8	
Interrupted aortic arch	2	0	
Single ventricle (DILV, TA, HLHS, etc.)	36	34	
Pulmonary atresia	13	11	
Transposition of the great arteries (d-, l- or congenitally corrected)	8	25	
Truncus arteriosus	2	3	
Criss cross heart, heterotaxy	3	2	
Eisenmenger	0	0	
Double outlet left ventricle	0	0	

Note, some admissions may have more than one cardiac diagnosis, so the total number of diagnoses exceeds the total for each group

Data from the Vizient Clinical Data Base used by permission of Vizient. All rights reserved *DILV* double inlet left ventricle; *HLHS* hypoplastic left heart syndrome; *TA* tricuspid atresia

with moderate and severe CHD who are hospitalized for COVID-19.

While young age is thought to be protective from COVID-19 for the general population, our findings do not support this for children or adults with CHD. In addition, the longer LOS and higher costs for patients with CHD is consistent with their expected fragility due to their underlying cardiac diagnoses, which is also highlighted by the higher severity of illness and mortality risk in the Pediatric group. It is also interesting that the adults with CHD had lower severity of illness at admission, but a similar mortality risk. The higher costs may also be due to more aggressive treatment provided to this population. There may be reductions in severe COVID-19 and hospitalizations with the introduction of vaccines against the SARS-CoV-2 virus, but this will not likely be available for young children for some time.

There are limitations to administrative database studies due to a lack of detail for some of the heart conditions and current functional status as well as potential errors in data entry. The code for COVID-19 (U07.1) is utilized for both symptomatic and asymptomatic individuals who have tested positive for the virus. However, it is unlikely that the current findings of worse outcomes in the CHD group were simply due to the group without CHD only having asymptomatic COVID-19 and the group with CHD only having symptomatic COVID-19. Despite these limitations, given the novelty of the COVID-19 pandemic and the relative rarity of moderate and severe CHD, using a large administrative database allows us to have a better sense of the current LOS (d)

Death (n, %)

Complications (n, %)

0.089

0.916 0.528

Pediatric (<18 years)	Cyanotic	Acyanotic	p
	(n = 109)	(n=261)	
LOS (d)	$28.0 \pm 41.3$	$34.9 \pm 57.0$	0.253
Complications (n, %)	9 (8.3)	34 (13)	0.192
Death (n, %)	3 (2.8)	15 (5.7)	0.222
Adult (≥18 years)	Cyanotic $(n=122)$	Acyanotic $(n = 388)$	р

Table 4 Hospital outcomes for pediatric (<18 years old) and adult ( $\geq$ 18 years) admissions for COVID-19 infection with cyanotic and acyanotic moderate or severe congenital heart disease

Data are presented as n (%) or mean ± standard deviation

Note, some patients had multiple diagnoses. Data from the Vizient Clinical Data Base used by permission of Vizient. All rights reserved *LOS* length of stay

 $9.9 \pm 11.3$ 

10 (8.2)

13 (10.7)

**Table 5** Severity of illness at admission and mortality risk for pediatric (<18 years) and adult ( $\geq$ 18 years) admissions for COVID-19 infection with and without congenital heart disease

Pediatric (<18 years)	$\begin{array}{c} \text{CHD} \\ (n = 160) \end{array}$	No CHD $(n=9318)$	р
	( <i>n</i> = 100)	( <i>n</i> =7516)	
Severity of illness			
Low/normal $(n, \%)$	128 (80)	8982 (96.4)	< 0.001
High ( <i>n</i> , %)	32 (20)	336 (3.6)	
Mortality risk			
Low/normal $(n, \%)$	77 (48.1)	6241 (67)	< 0.001
High ( <i>n</i> , %)	83 (51.9)	3,077 (33)	
Adult (>18 years)	CHD ( <i>n</i> = 389)	No CHD ( <i>n</i> =657,841)	р
Severity of illness			
Low/normal $(n, \%)$	174 (44.7)	235,795 (35.8)	< 0.001
High ( <i>n</i> , %)	215 (55.3)	422,046 (64.2)	
Mortality risk			
Low/normal $(n, \%)$	307 (78.9)	526,085 (80)	0.605
High ( <i>n</i> , %)	82 (21.1)	131,756 (20)	

Data are presented as n, %. Data from the Vizient Clinical Data Base used by permission of Vizient. All rights reserved

status of the COVID-19 pandemic than any single center study could provide.

The current study suggests worse hospital outcomes when patients with CHD are hospitalized for COVID-19 infection, including higher mortality for children. These data stress the importance of primary prevention with vaccination, social distancing and masking measures to reduce severe COVID-19 and hospitalizations and also to increase herd immunity to protect the children who are too young to receive the vaccines at this time. These findings can help to further guide treatment strategies and prioritize patients for vaccination. Acknowledgements None.

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 $12.6 \pm 16.3$ 

33 (8.5)

34 (8.8)

Data Availability Data not available.

Code Availability N/A.

## Declarations

**Conflict of interest** All authors declare that they have no conflict of interest.

**Ethical Approval** All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional and/or national research committee and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards.

**Informed Consent** The need for informed consent was waived for this study of deidentified data.

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