


# Factors affecting COVID-19 outcomes in patients with congenital heart disease

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## Letter to the Editor

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Haiduc et al attempt to summarise the current literature surrounding the effect of congenital heart disease (CHD) on COVID-19 outcomes.<sup>1</sup> We propose further consideration of the following factors: age, type and severity of CHD, and comorbidities.

This review directly compares studies and case reports of patients with ages ranging from 3 months to 76 years. Discrepancy in ages is important to consider since COVID-19 is known to affect infants less than 1 year of age more severely.<sup>2</sup> This may be due to low T-cell activation, reduced expression of ACE2 in the lungs and other developmental differences.<sup>2</sup> Older patients are likely to have developed more comorbidities which may be the reason for the presentation of worse COVID-19 symptoms, rather than being due to their CHD alone. Literature also shows racial disparities in the severity of COVID-19 which may confound the effect of CHD on outcomes.<sup>3</sup>

Contrary to this review, there is evidence that individual defects such as single ventricular defects are not sufficient cause alone for worse COVID-19 outcomes.<sup>4</sup> Lewis et al reported that out of 53 CHD patients, there appeared to be no correlation between the complexity of the CHD and the subsequent infection-related cardiac decompensation displayed by the patient.<sup>4</sup> This may explain why patients with similar CHDs experience different disease trajectories. Therefore, using only the CHD status is not a sufficient indicator for poor outcomes and markers of symptom progression such as raised cardiac troponins upon admission may prove more useful.<sup>5</sup> This is particularly relevant since CHD patients with infection-induced myocardial injury are at a higher risk of complications.<sup>5</sup> Treatment protocols used to manage infections should also be considered amongst the studies included in the review. For example, Non-steroidal anti-inflammatory drugs use in early infection is thought to be controversial due to reports of causing more complications when administered for respiratory tract infections.<sup>6</sup>

From the studies included in the review, 24% of the patients had comorbidities with the most frequent being genetic conditions, type 2 diabetes mellitus, and chronic kidney disease.<sup>1</sup> Massin et al demonstrates that a significant proportion of children with CHD have associated non-cardiac comorbidities including genetic conditions, with the most common being trisomy 18.<sup>7</sup> These comorbidities make it difficult to assess the direct effect of CHDs on outcomes. Similarly, Agarwal et al show that adults with CHD are twice as likely to have non-cardiac comorbidities compared to their non-CHD counterparts.<sup>8</sup> Additionally, for patients below the age of 40, the more severe the CHD, the higher the risk of developing non-cardiac comorbidities.<sup>8</sup>

In conclusion, in order to determine causality or a significant correlation between COVID-19 prognosis and CHD, this review needs to stratify CHD according to type of defect and severity, whilst controlling for confounders such as age and comorbidities.

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