

# The Correlation Between Congenital Heart Disease and The Risk of ADHD in Children

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# ABSTRACT

Congenital Heart Disease (CHD) is the most common birth defect in the world, and patients are more prone to intellectual functioning deficits, developmental issues, and academic performance issues. Attention Deficit Hyperactivity Disorder (ADHD) is a neurodevelopmental disorder and is associated with coronary heart disease (CHD). While cardiac patients with ADHD symptoms are frequently misdiagnosed and, as a result, undertreated, this study examines the association between CHD and the risk of ADHD, including the effect of cardiac surgery on children. The data for this study was compiled by searching multiple databases using pertinent keywords. Through school age and adolescence, children with CHD have an elevated risk of neurodevelopmental impairment, specifically ADHD. In children with surgically corrected ADHD, inattention becomes the predominant subtype, and scores were significantly increased. However, the severity of the symptoms is reduced if cardiac surgery is performed at a young age. This association leads to alterations in behavior associated with hypoxic-ischemic mechanisms that occur in oxygen-sensitive regions. To improve the quality of life of children with CHD, surveillance, screening, evaluation, and management strategies for ADHD should be promoted regardless of their specific cardiac diagnosis.

Keywords: congenital heart disease; attention deficit/hyperactivity disorder; children; cardiac surgery

## INTRODUCTION

Congenital heart disease (CHD) is the most prevalent birth defect in the world, affecting millions of infants each year (1). Between 1970 and 2017, the global mean prevalence of CHD was 8.22 per 1000. Globally, the prevalence of CHD increased by 10% every five years (2). Children with CHD are more susceptible to intellectual functioning deficits, developmental problems, and academic performance issues (3). Children with congenital heart disease are at increased risk for several cognitive disorders, including inattention, which can interfere with school-based learning (2). Previous studies have reported unfavorable effects of chronic and intermittent hypoxia on development, behavior, and academic achievement in children with CHD.

It has been reported that children with congenital heart disease have a 30% higher risk of inattention and hyperactivity disorder compared to healthy individuals. Importantly, despite the fact that nearly half of surgically treated patients require remedial school services when they reach adolescence, cardiac patients with ADHD symptoms are frequently underdiagnosed and, as a result, undertreated (5).

Attention deficit hyperactivity disorder (ADHD) is an important neurodevelopmental outcome because it is a treatable disorder.

The morbidity associated with undiagnosed and untreated ADHD is substantial (6). Hansen et al. found that DSM-IV estimated the prevalence of ADHD to be between 3% and 5% (7). ADHD is also associated with maternal anxiety, but it is advantageous to administer ADHD medication if its severity is detected early (8). There are numerous screening instruments available for detecting ADHD. The Swanson, Nolan, and Pelham IV (SNAP-IV) Teacher and Parent Rating Scale is a 20-item questionnaire based on the Diagnostic and Statistical Manual of Mental Disorders, fourth revision diagnostic criteria for ADHD.

Hansen et al. estimated a correlation of 41.2% between CHD and ADHD parent ratings based on the SNAP-IV (7). Consequently, the purpose of this study is to examine the association between CHD and the risk of ADHD, as well as the effect of pediatric cardiac surgery. Through our review of the literature, we hope to improve the quality of life for children with CHD and ADHD.

### METHOD

The correlation between congenital heart disease and the risk of childhood attention deficit hyperactivity disorder (ADHD) was the subject of this literature review, which was compiled using data from PubMed, Science Direct, Research Gate, and other databases.

The final article is from April 2022. The reviewed literature is written in English or Indonesian for the authors' convenience.

No.	Author (Year)	Country	Sample Characteristics			Outcome Data	
			Age (Mean ± SD)	Gender (Boy %)	History of cardiac surgery	ADHD Prevalence	Cardiac surgery
1	Czobor (2021)	Hungary	School aged (10.5 ± 3.3)	51%	54 children having the first surgery under 3 years of age and 26 patients received the first operation at or above 3 years of age	Those undergoing surgery later in life had higher prevalence rates than the other two subgroups.	There were statistically significant differences in the length of hospital stay, the number of operations, and the frequency of perioperative cyanosis; values were higher for children who were first operated below 3 years of age.
2	Razzaghi (2015)	USA	0 - 17	49,4%	-	Children 2–17 years of age, children with CHD diagnoses had higher odds of ADHD.	÷
3	Tsao (2017)	Taiwan	<18 (2.21 ± 4.08)	48,9%	<b></b>	Incidence of ADHD were higher in the CHD group than in the control group.	
4	DeMaso (2017)	USA	10-19 (14.5 ± 3.0)	61%	All pateints underwent fontan procedure	Patients were more likely to be diagnosed with lifetime disruptive behavior disorders, specifically ADHD (34%)	-
5	Yamada (2013)	Canada	7-15		All children underwent open-heart surgery before 1 year of age	The prevalence of a positive screening score for ADHD in subjects who had undergone early cardiac surgery was much higher at 29% than in healthy control group with 3,3%.	Children who have open-heart surgery at younger than 1 year of age for CHD are more likely than healthy control subjects to have a SNAP-IV score suggestive of ADHD.
6	Hansen (2012)	USA	7-15 (12.6)	63%	÷.	Patients with CHD were reported to have elevated SNAP-IV scores by parents and counselors (11.8%).	
7	Holst (2020)	Denmark	10-16 (13 ± 1.8)	55%	All children went cardiac surgery	Children with surgically corrected CHD had significantly increased inattention- and total attention deficit/hyperactivity disorder symptom scores compared with control.	In children who undergo CHD surgery later than 2 weeks after birth, it has been shown they have impaired brain growth and delayed development of language skills compared with children who undergo surgery during the first 2 weeks of life.
8	Shahri (2021)	Iran	${}^{<18}_{-45.84\ months)}$	78.70%	22,1% underwent surgery	The prevalence of ADHD in CHD population was high (31.6%).	-
9	Loblein (2022)	England	3-21 (8.6)	56,8%	-	27.5% of patients were diagnosed with ADHD, which is significantly higher than the prevalence in the general population (9.04%).	-
10	Gonzalez (2021)	USA	4-17	56,8%	-	A prevalence of ADHD in youth with CHD aged 4 to 17 years of 5.1%, whereas the non- CHD population had a lower prevalence of ADHD of 2.1%.	υ.
11	Wang (2020)	Taiwan	6-15 (10.2 ± 2.43)	53,5%	57,8% patients went surgery with age at frst surgery (years, mean, range) 1.3 (0.1– 12.0)	The overall prevalence of ADHD was 12.4%, including 3.2% for the combined subtype, 6.8% for the inattentive-predominant subtype, and 2.4% for the hyperactivity/impulsive-predominant subtype.	The highest prevalence of the inattention-predominant subtype of ADHD was observed in the children with cyanotic CHD, in those who received cardiac surgery or cardiopulmonary bypass, and in those with a high disease severity index.
12	Shillingford (2008)	USA	5.4-10.4 (7.9 ± 1.5)	67%	Each patient underwent cardiac surgery before 45 weeks postconception age with planned DHCA and CPB.	5% would be highly likely to have ADHD, and another 15% would be considered at risk for having a diagnosis of ADHD.	There were no statistically significant differences between the children with a Fontan operation for single ventricle and those who underwent biventricular repair, nor were there associations between adverse outcomes and the duration of intraoperative support, such as CPB and DHCA.
13	Gaudet (2021)	Canada	$5\ (5.55 \pm \\ 0.26)$	52,7%	All patients were having undergone at least 1 invasive procedure.	Prevalence ADHD or ADHD-related symptoms is much higher in children with CHD.	-

### **TABLE 1 :** Studies included to this review.

### **OVERVIEW**

Congenital Heart Disease (CHD) is a heart defect present at birth. Congenital heart disease (CHD) is also referred to as congenital heart defects, which is a general term for abnormalities in the structure of the heart and great blood vessels that are present at birth, are frequently observed, and are the leading cause of death among all types of congenital abnormalities (9). Due to cardiac abnormalities, hemodynamic disturbances can produce symptoms that characterize the severity of the abnormality. The presence of growth retardation, cyanosis, decreased exercise tolerance, frequent recurrent respiratory tract infections, and the ability to hear heart sounds may be early indicators of heart defects in infants and children (10).

Two possible causes of congenital heart disease are genetic and environmental factors. Genetic factors include heredity or a family history of disease, as well as Down syndrome and other syndromes caused by an abnormal number of chromosomes. Maternal rubella virus infection, use of teratogenic drugs during pregnancy, and excessive alcohol consumption are environmental factors (11).

Important basic investigations for CHD include chest xrays, electrocardiography, and routine laboratory tests, whereas important follow-up examinations include echocardiography and a cardiac catheterization procedure. The combination of two advanced examinations for visualization and confirmation of the morphology and path-anatomy of each type of congenital heart disease allows for a near-perfect diagnosis (12).

In general, the treatment of congenital heart disease includes both non-surgical and surgical methods. Nonsurgical management includes medical management and interventional cardiology. Generally, medical management is secondary due to complications from the heart disease or other accompanying disorders (13).

Congenital Heart Disease (CHD) is the most common birth defect in the world, and patients are more prone to intellectual functioning deficits, developmental issues, and academic performance issues.

Attention Deficit Hyperactivity Disorder (ADHD) is a neurodevelopmental disorder and is associated with coronary heart disease (CHD). While cardiac patients with ADHD symptoms are frequently misdiagnosed and, as a result, undertreated, this study examines the association between CHD and the risk of ADHD, including the effect of cardiac surgery on children.

The data for this study was compiled by searching multiple databases using pertinent keywords. Through school age and adolescence, children with CHD have an elevated risk of neurodevelopmental impairment, specifically ADHD. In children with surgically corrected ADHD, inattention becomes the predominant subtype, and scores were significantly increased. However, the severity of the symptoms is reduced if cardiac surgery is performed at a young age. This association leads to alterations in behavior associated with hypoxicischemic mechanisms that occur in oxygen-sensitive regions.

To improve the quality of life of children with CHD, the surveillance, screening, evaluation, and management strategy for ADHD should be promoted regardless of their specific cardiac diagnosis.

### Attention Deficit/Hyperactivity Disorder (ADHD)

Multiple tests are utilized to diagnose children with ADHD. Additionally, sleep disturbances, anxiety, depression, and learning disorders all share the same symptoms. ADHD is diagnosed using the Diagnostic and Statistical Manual, Fifth Edition (DSM-5) from the American Psychiatric Association. DSM-5 is applicable from early childhood to adulthood. Patients with ADHD exhibited persistent patterns of inattention and hyperactivity (14).

In Indonesia, there are two screening instruments for ADHD. Parents and teachers can complete both the Skala Penilaian Perilaku Anak Hiperaktid Indonesia (SPPAHI) and the Abbreviated Conner's Teacher Rating Scale (ACTRS). If the score exceeds the threshold, the child can be identified as a child at high risk (14). Children with ADHD are more likely to exhibit oppositional behavior, academic failure, family conflict, forgetfulness, impulsive behavior, difficulty adapting, and poor learning achievement. These accomplishments require psychomotor (activity), cognitive (knowledge), and behavioral (emotional) components (15).

In school-aged children with dextro-transposition of the great arteries (D-TGAs), deficits in executive functions (EF) have been reported. On the Behavior Rating Inventory of Executive Functions (BRIEF), parents and educators of children with congenital heart disease (CHD) demonstrated significant difficulties, particularly in working memory. Calderon et al. also reported EF deficits in preschool and school-aged children (16).

# Current Evidence on the Association of CHD and ADHD

On the basis of a literature search in multiple databases using pertinent keywords, fourteen works of literature that met the inclusion and exclusion criteria were identified. The majority of studies utilized school-aged participants. As the symptoms of ADHD become more apparent in patients, it becomes possible to assess and establish the diagnosis. Six studies used a sample of all CHD patients who underwent cardiac surgery, and the majority of them were young.

Comparing the prevalence, incidence, or odds of CHD in children to those of healthy controls, the vast majority of the studies pooled for this literature review indicate a statistically significant increase. A large cohort study conducted in Taiwan gives results of the overall prevalence of probable ADHD in all patients with CHD was 12.4%. There is a significantly higher prevalence of the inattention-predominant subtype among CHD patients, particularly in patients with cyanotic heart disease and those with a history of attention deficit hyperactivity disorder has undergone cardiac surgery (17). However, the predominant subtypes of hyperactivity/impulsivity did not differ significantly (17). Several studies have yielded dissimilar prevalence results. The prevalence of subjects with a positive screening score for ADHD was much higher in the CHD group, at 29%, according to a Canadian cohort study (18). The prevalence of ADHD in the CHD population was found to be 31.6% (2). In the most recent study, conducted by Loblein et al., the prevalence of patients diagnosed with ADHD was significantly higher than in the general population (9.04%) (19).

According to a previous cross-sectional study (7), a significant proportion of children with CHD exhibited symptoms of ADHD. This association led to behavioral changes associated with hypoxic-ischemic damage to the brain's white matter (20). Patients born with CHD are exposed to physiological stressors that continue to evolve as they mature. One of the stressors is the level and duration of hypoxemia that occur in highly oxygensensitive regions, which are the prefrontal cortex and corpus striatum of the brain. This region is presumed to be connected to the executive control network of attention (7; 20). CHD patients have risk factors for the inattention-predominant subtype, which include postoperative seizure and a number of cardiopulmonary bypass surgeries, as the greatest risk, cyanotic CHD, prior use of ECMO, and male sex (17). Preoperative hypoxemia in infants with cyanotic CHD is associated with a higher risk of attention deficits compared to acyanotic CHD and healthy infants (20).

In the high severity index, the inattention-predominant subtype of ADHD has the highest prevalence. However, there was no difference in the prevalence of hyperactivity/impulsivity-related symptom severity (17; 7). This high prevalence was observed in children with cyanotic CHD, those who underwent cardiac procedures, and those with an elevated disease severity index (17). ADHD symptoms were more severe in CHD patients treated surgically at or above 3 years of age compared to both the control group and those treated surgically at a younger age (21). Consistent with the findings of other studies, children with Transposition of the Great Arteries who undergo CHD surgery more than two weeks after birth exhibit impaired brain growth and delayed language development compared to children treated within the first two weeks (20).

Anxiety, depression, and post-traumatic stress were frequently observed with significantly elevated scores in children with CHD and ADHD (22; 23). Achievements in school, education, or work functioning can be disrupted in ADHD patients. Recent research (20) indicates that children with CHD and additional ADHD have a lower quality of life score. Then, neurodevelopmental disorders, particularly ADHD, should be evaluated in all children at high risk for CHD (19).

### CONCLUSION

Through school age and adolescence, children with congenital heart disease (CHD) have an increased risk of neurodevelopmental impairment, particularly Attention Deficit Hyperactivity Disorder (ADHD). On the basis of prevalence rates, inattention symptoms become the predominant type of ADHD and are significantly more prevalent hypoxia-induced ischemia. The ADHD scores of surgically corrected CHD children were significantly higher. However, the symptoms are less severe if cardiac surgery is performed at a young age.

To improve the quality of life of children with CHD, the surveillance, screening, evaluation, and management strategy for ADHD should be promoted regardless of their specific cardiac diagnosis. Other neurocognitive and psychiatric evaluations are advised throughout a child's development.

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### **Conflict of interest**

The authors declared there to be no conflict of interest in this study.

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### Author contribution

YS carried out the idea of this study and revision. NPN, ARHA, and RDAJH contribute to the design of the study, interpreted the results, and arrangement of the manuscript.

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