

# LJMU Research Online

Lotto, R and Lotto, A

Delayed Sternal closure following complex cardiac surgery in neonates https://researchonline.ljmu.ac.uk/id/eprint/26686/

#### **Article**

**Citation** (please note it is advisable to refer to the publisher's version if you intend to cite from this work)

Lotto, R ORCID logoORCID: https://orcid.org/0000-0002-4305-0513 and Lotto, A Delayed Sternal closure following complex cardiac surgery in neonates. World Journal for Pediatric and Congenital Heart Surgery. ISSN 2150-1351 (Accepted)

LJMU has developed LJMU Research Online for users to access the research output of the University more effectively. Copyright © and Moral Rights for the papers on this site are retained by the individual authors and/or other copyright owners. Users may download and/or print one copy of any article(s) in LJMU Research Online to facilitate their private study or for non-commercial research. You may not engage in further distribution of the material or use it for any profit-making activities or any commercial gain.

The version presented here may differ from the published version or from the version of the record. Please see the repository URL above for details on accessing the published version and note that access may require a subscription.

For more information please contact <a href="mailto:researchonline@ljmu.ac.uk">researchonline@ljmu.ac.uk</a>

# Delayed Sternal closure following complex cardiac surgery in

2	neonates
3	
4 5	Amr Ashry <sup>1,2</sup> *, Shubhi Gupta <sup>3,4</sup> *, Hesham Hassan Ahmed <sup>5</sup> , Robyn Lotto <sup>6,7</sup> , Ramesh Kutty <sup>1</sup> , Rafael Guerrero <sup>1,4</sup> , Ram Dhannapuneni <sup>1</sup> , Attilio A. Lotto <sup>1,6,7</sup>
6 7	1. Department of Paediatric Cardiac Surgery, Alder Hey Children's Hospital, Liverpool, United Kingdom.
8 9	2. Department of Cardiothoracic Surgery, Assiut University Hospital, Faculty of Medicine, Assiut University, Assiut, Egypt.
LO	3. Department of Surgery, Aintree University Hospital, Liverpool, UK.
l1	4. School of Medicine, University of Liverpool, Liverpool, UK.
L2	5. Department of Cardiothoracic Surgery, Faculty of Medicine, Menoufia University, Egypt.
L3	6. Faculty of Health, Liverpool John Moores University, Liverpool, United Kingdom.
L4	7. Liverpool Centre for Cardiovascular Health, Liverpool, United Kingdom.
L5	
L6	Corresponding Author:
L7	Mr. Attilio Lotto, MD, MCh, FRCS-CTh
L8	Consultant in Congenital Cardiac Surgery &ECMO Service Co-Lead
L9	Alder Hey Children's Hospital
20 21	Professor in Congenital Cardiac Surgery at Liverpool John Moores University & Liverpool Centre for Cardiovascular Science
22	E-mail: attilio.lotto@alderhey.nhs.uk
23	
24	We presented our results in Porto, Portugal at AEPC 2024 Annual Meeting.
25	Key words: Delayed sternal closure, congenital cardiac surgery, sternal wound infection,
26	neonatal cardiac surgery.
27	*: Authors AA & SG had equal contribution to the manuscript.
28	
29	Word count: 3851 words

#### Abstract

- 3 Background: Delayed sternal closure (DSC) is a well-established surgical intervention
- 4 following complex congenital cardiac surgeries mitigating postoperative haemodynamic and
- 5 respiratory instability. It is mostly used in neonates requiring prolonged cardiopulmonary
- 6 bypass, aortic cross-clamp times or deep hypothermic circulatory arrest, predisposing to
- 7 myocardial oedema or bleeding. Our study evaluates morbidity and mortality after DSC in
- 8 neonates including superficial, deep sternal wound infections and requirement of surgical
- 9 debridement.
- 10 Methods: Retrospective review of neonates who underwent DSC after cardiac surgery in a
- single centre from 2015 to 2021.
- Results: 187 neonates were identified. Mean age and weight were  $12.8 \pm 6.8$  days and  $3.3 \pm$
- 13 0.5 kg, respectively. Mean days of opened chest were  $3.8 \pm 5.8$  days. Two neonates (1.07%)
- required sternal wound debridement, whilst 19 cases (10.2%) had superficial wound infections.
- Mean ICU and hospital stay were  $12.8 \pm 16.6$  and  $25.9 \pm 36.9$  days. 30-days mortality occurred
- in 9 cases (4.8%). Univariate analysis indicated that DSC days (p= 0.01), ECMO (p= 0.000),
- aortic cross clamp time (p= 0.007) and CPB time (p=0.006) to be associated with 30-days
- mortality, whilst in multivariable analysis, only ECMO was significant (p= 0.002). RACHS-1
- score was the only independent risk factor for sternal wound infection in univariate analysis (p
- =0.019) and multivariable analysis (p=0.05).
- 21 Conclusion: DSC is a safe therapeutic option following complex neonatal cardiac surgery,
- 22 where cardiac compression by sternal approximation is not tolerated because of, myocardial
- oedema, haemodynamic instability or coagulopathy. Higher RACHS-1 score was associated
- 24 with greater incidence of sternal wound infections.

#### Introduction

- 3 Delayed sternal closure (DSC) is a widely recognised surgical intervention aimed at mitigating
- 4 postoperative haemodynamic and respiratory instability in paediatric patients following
- 5 complex cardiac surgery [1]. Approximately 10% of paediatric cardiac patients leave the
- 6 operating room with an open chest, with subsequent sternal closure performed after an average
- 7 duration of three days [2–4].
- 8 It is most frequently indicated in complex cases involving prolonged cardiopulmonary bypass
- 9 and aortic cross-clamping, which can potentially lead to intraoperative bleeding and
- myocardial oedema [5]. In such circumstances, sternal closure can trigger haemodynamic
- instability with reduction in cardiac output and diastolic filling [5]. Delayed chest closure can
- improve morbidity and mortality for patients following complex paediatric cardiac operations,
- by preventing the onset of haemodynamic instability.
- 14 However, DSC may predispose patients to sternal wound infections (SWI) and mediastinitis
- 15 [6-9]. SWIs can be categorised into two types: superficial SWIs, managed conservatively with
- antibiotics, and deep SWIs which require surgical debridement. Albeit rare, severe
- complications such as sepsis, can be life-threatening [6]. Moreover, mortality rates in this
- critically ill patient subgroup vary considerably, ranging from 8 % to as high as 34% [1-6].
- 19 Considering the pros and cons of DSC, a lack of consensus exists within the literature
- 20 concerning the indication for or timing of sternal closure in these patients. Moreover, different
- 21 clinical series report variable morbidity and mortality rates associated with this procedure. A
- prevalence of 3.5-18.0 % for SWIs following delayed sternal closure has been documented in
- 23 the literature [2-4, 6-15]. This retrospective review aims to evaluate the morbidity and
- 24 mortality following delayed sternal closure in the neonatal population, focusing on the

1 occurrence of superficial and deep sternal wound infections and the requirement of surgical

2 debridement.

3

4

5

7

8

9

10

12

13

14

15

16

#### Material and methods

# Study design and patient population

6 This study has been approved by Liverpool John Moores University Ethical committee (UREC

reference number: 25/NAP/001). A retrospective data collection was performed for neonates

who underwent DSC following congenital cardiac surgery via sternotomy from 2015 to 2021,

at Alder Hey Children's Hospital, Liverpool, United Kingdom. Patients who are over 28 days

of age and patients with incomplete data were excluded.

11 Variables including indication of DSC, time to sternal closure, mean paediatric intensive care

unit (PICU), hospital stay duration, superficial & deep wound infections, 30-day mortality,

sepsis and surgical risk according to the Risk Adjustment for Congenital Heart Surgery

(RACHS-1) score, need for extracorporeal membrane oxygenation (ECMO) were extracted

from our database. Data relating to incidence of SWI, surgical intervention and 30-day

mortality were double checked by another author.

17

18

20

21

22

23

### **Endpoints**

19 The primary endpoints of this study were the superficial and deep sternal wound infection and

mediastinitis rates with the need for sternal wound debridement rate. Superficial SWIs were

defined as inflammatory changes confirmed with wound swabs. Deep SWIs were confirmed if

the muscular layers were affected. Mediastinitis is defined as a positive pathogen identified by

culture swabs affecting the mediastinum and the sternum. Secondary endpoints involved

- 1 evaluating associated morbidity such as sepsis, intensive care and hospital stay duration and
- 2 30-day mortality rates.

4

5

7

8

9

10

11

12

13

14

15

16

17

18

19

20

#### **Indication for DSC**

6 The decision to delay sternal closure is determined preoperatively for operations like the

Arterial Switch Operation for transposition of the great arteries and the Norwood procedure for

hypoplastic left heart syndrome. In other instances, it is decided intraoperatively in case of

myocardial oedema and myocardial distention from a variety of clinical conditions like low

cardiac output, arrhythmias, respiratory failure needing high ventilation pressures.

Coagulopathy with the need to leave the chest packed with swabs is a further indication or in

patients with postoperative circulatory support through central ECMO. Also, we adopt the DSC

policy in the case of pulmonary artery banding (PAB) for single ventricle circulation or in case

of septal defects (Multiple VSDs, Apical VSD or complete AVSD not closable in conjunction

with hypoplastic aortic arch repair under circulatory arrest), as we have experienced, in such

cases, that the PAB will need further adjustments few days after surgery. We use intravenous

(IV) Cefuroxime (Cephalosporin) during the days of opened chest, and we tend to continue it

for 2 doses after chest closure. In case of cephalosporin or penicillin allergy, we use IV

Teicoplanin plus IV gentamicin as an alternative to cefuroxime and we continue them for 2

doses after chest closure.

21

22

#### Surgical techniques:

- 2 Our approach to open chest treatment encompasses various techniques depending on the
- 3 surgeon's preference. In cases who went out of theatre on ECMO, we utilise silastic membrane
- 4 that is sutured to the skin with continuous 5/0 polypropylene sutures, then we further cover it
- 5 with 2 layers of antimicrobial drape (Ioban 3M).
- 6 In non-ECMO cases, we utilise different techniques based on the individual surgeon's
- 7 preference which can be:
- 8 1- silastic membrane that is sutured to the skin with continuous 5/0 polypropylene sutures,
- 9 then we further cover it with 2 layers of antimicrobial drape (Ioban 3M).
- 10 2- Direct skin closure using subcuticular Monocryl sutures, leaving the sternal edges open.
- 11 Then, covering the wound with sterile dressing.
- 12 3- Sterile dressings over a small size swab left in the chest.
- 4- Antimicrobial drape (Ioban 3M) alone over the open chest with 1 small swab left inside
- 14 the chest.
- 15 5- Few cases needed the use of sternal bridge to keep the sternal edges apart with a piece of
- 16 chest drain tube sutured to the sternal edges to prevent compression to the heart or RV-PA
- 17 conduit. Then a silastic membrane is sutured to the skin edges covered by a sterile dressing.
- 18 Upon chest closure, any dressing is removed in a sterile fashion, to expose the sternum
- 19 removing retained swabs, mediastinal cavity washout with warm saline solution, followed by
- standard sternotomy closure in layers using 2/0 PDS for the sternum, 3/0 Vicryl for the muscle
- 21 and subcutaneous layers, and 5/0 Monocryl for subcuticular skin closure. We do all our chest
- 22 closures in ICU including those who need ECMO weaning and decannulation. We do not use

- 1 GORE-TEX membrane to cover the heart routinely. Its use is a surgeon's preference. In case
- 2 of needing to cover the heart or great vessels, suturing a ePTFE 0.1 mm membrane (Gore-Tex)
- 3 to the pericardial edges with polypropylene 5/0 interrupted sutures is done at the time of chest
- 4 closure in ICU.

## Regarding the cases who had sternal wound infection:

- 6 Out of the two cases who required wound revision and debridement in the operation room, the
- 7 first patient exhibited sternal dehiscence on examination and had positive mediastinal wound
- 8 swab cultures. The second patient developed signs of a deep sternal wound infection,
- 9 characterised by a high C-reactive protein, pyrexia, positive blood cultures and chest swab
- 10 cultures, as well as purulent material on swabs covering the midline sternotomy incision with
- 11 sternal dehiscence.
- 12 However, in cases of superficial wound infection, we managed it conservatively with
- antibiotics based on the culture and sensitivity results and regular dressings. If the baby needed
- 14 frequent dressing change because of superficial discharge, we tend to use PICO negative
- pressure wound dressing and we change it every 5-7 days.

# Statistical analysis

- 17 Categorical data was reported as frequencies and proportions. Mean and standard deviation
- were used to express quantitative data. Univariate and multivariable regression analyses were
- conducted to identify risk factors for sternal wound infection and 30-day mortality. The level
- of significance was considered at p value < 0.05. Statistical analyses were conducted with
- 21 SPSS Version 27.0 (SBSS Inc, Chicago, IL).

22

16

#### Results

1

2

Demographic data

Over a 6-year period, 1896 patients were identified and screened of which 187 neonates were 3 included in our patient cohort. Mean age and weight at operation were  $12.8 \pm 6.8$  days and 3.34 5  $\pm$  0.5 kg, respectively. Mean aortic cross clamp time and CPB time were 102.8  $\pm$  52.5 and 183 6  $\pm$  81 minutes, respectively. Mean RACHS-1 Score was 4  $\pm$  1.05. Arterial Switch Operation, with or without Ventricular Septal Defect (VSD) repair 70 (37.4%), Hypoplastic Aortic Arch 7 Repair (HAA) 49 (26.2%), Norwood operation 30 (16.0%) comprised the most frequent 8 operations performed in the DSC cohort as depicted in Figure 1. 9 Mean days of open chest were  $3.8 \pm 5.8$  days. Only 2 patients (1.07%) needed sternal wound 10 11 debridement in theatre for deep sternal wound infections, whilst 19 cases (10.2%) had superficial wound infections managed conservatively with antibiotics. Mean days of wound 12 infection was  $10.76 \pm 6.65$  days and the 2 cases who needed wound revision and debridement 13 was at the 7th and 17th day after chest closure. 14 15 Mean ICU and hospital stay were  $12.8 \pm 16.6$  and  $25.9 \pm 36.9$  days respectively. Pre-operative ECMO was required in 1 neonate (0.5%). Intra-operative ECMO was required in 6 cases 16 (3.2%), whilst 21 patients (11.2%) needed post-operative ECMO. 52 cases (27.8%) required 17 further chest explorations. 30-day hospital mortality occurred in 9 cases (4.8%). 24 neonates 18 (12.8%) developed sepsis within 30 days of primary operation. Demographic data has been 19 summarised in Table 1. 20

21

22

23

#### Regression analysis

1

- 2 Univariate regression analysis showed that the RACHS-1 score (p=0.019, 95% CI -0.047-
- 3 0.178) was found to be significantly associated with sternal wound infections as described in
- 4 Table 2. Furthermore, DSC days (p=0.011, 95% CI -0.19 -0.003), ECMO (p =0.000, 95% CI
- 5 0.251-0.465), a ortic cross clamp time (p=0.007, 95% CI 0.000- 0.003) and CPB time (p=0.006,
- 6 95% CI -0.002-0.000) were found to be significantly associated with 30-days mortality, in the
- 7 univariate regression analysis, which has been depicted in Table 3.
- 8 On multivariate analysis, the RACHS-1 score was found to be the one significant independent
- 9 risk factor for sternal wound infections, p= 0.05, 95% CI 0.995-4.012 as described in Table 4.
- 10 In addition, ECMO was found to be the sole significant risk factor for 30-day mortality,
- 11 (p=0.002, 95% CI 0.000-0.122). Multivariate analysis for 30-day mortality is shown in Table
- 12 5.

13

14

# Discussion

15 Median sternotomy with delayed sternal closure has become standard practice in selected subgroup of patients needing to optimise postoperative recovery. This strategy proves 16 particularly advantageous in case of haemodynamic or respiratory instability, myocardial 17 18 oedema and dilation, when coagulopathy and bleeding require packing of the chest or the necessity for central ECMO support. For some subgroup of patients, the decision of leaving 19 the chest open is taken preoperatively, like in case of ASO for TGA, or following the Norwood 20 Procedure for HLHS. In other cases, when is likely that additional procedures need to be 21 performed, like tightening a PAB, the DSC is adopted. Adopting a DSC policy is not without 22 risks, for instance patients with prolonged CPB time, low cardiac output, or coagulopathy 23 needing transfusion are recognised be at higher risk for sternal wound infections [1]. 24

- 1 Furthermore, SWIs have been associated with longer postoperative stays, as well as increasing
- 2 the cost of stay due to the requirement for prolonged ventilation [6].
- 3 A significant challenge in comparing our findings with existing literature lies in the lack of
- 4 standardization in defining infection subtypes. Variability in the criteria used to classify
- 5 superficial and deep wound infections across studies introduces potential inconsistencies and
- 6 may account for some of the observed differences in reported incidence rates [6-16].
- 7 There exists a variance in the reported incidence rates of sternal wound infections in patients
- 8 with DSC. Some retrospective series have found superficial SWI rates as (6.7-9.7%) and deep
- 9 SWIs as (3.9-10.5%), respectively[1, 6, 16]. Whilst others have reported no significant
- association between DSC and an increase in surgical site infections. Von Stumm et al described
- 11 7.3% superficial SWI rate in a 358 paediatric cohort but no deep SWIs requiring surgical
- debridement[1]. Furthermore, Yang et al reported a 9.7% surgical site infection rate in neonates
- post DSC, of which there were 28 cases of mediastinitis[16]. Our retrospective study
- demonstrates similar superficial SWI rates, 10.2%, whilst our deep SWI rate is considerably
- 15 lower, 1.07%.
- Tabbutt and Özker et al reported a 19% mortality rate following paediatric DSC [6, 17].
- Hurtado-Sierra et al reported mortality as 22.4% with patients a RACHS- 1 score  $\geq$  3 posing a
- higher risk of mortality [3]. Our study shows a considerably lower 30-day mortality rate of
- 19 4.8%. 30-day mortality may be attributed to nosocomial infections. 12.8% of our neonates
- developed sepsis within 30 days of admission, significantly lower than Elassal et al, 54.1% in
- a paediatric cohort [18]. Infection rates are reported to be higher in patients with prolonged
- 22 ICU stay [18]. Our study depicts an association between prolonged ICU stay and the risk of
- SWIs on univariate analysis. Our ICU and hospital stay 12.8 (SD 16.6) and 25.9 (SD 36.9)

- 1 respectively is higher than other neonatal cohorts; Yang et al report a ICU and hospital length
- 2 of stay in their SWI as 11.64 (SD 2.6) and 22.26 (SD 4.3), respectively[16].

- 4 On univariate analysis, cardiopulmonary bypass time, prolonged DSC time and aortic cross
- 5 clamp time were significantly associated with 30-day mortality. Cardiac surgery-related
- 6 morbidity has been primarily attributed to the use of cardiopulmonary bypass. This can be
- 7 explained by the fact that extracorporeal circulation can induce a systemic inflammatory
- 8 response, trauma, ischemia and reperfusion injury, endothelial dysfunction, and activation of
- 9 the coagulation cascade [19-22]. Prolonged cross-clamp time has also been shown to be an
- independent predictor of 30-day mortality in the literature, similar to our findings [25].
- Furthermore, aortic cross clamp time of more than 60 mins with a 91.2% higher probability of
- sternal wound infections. Our study, on the other hand, did not found such association [16].

- Our retrospective study found one significant association between a higher RACHS-1 score
- and risk of SWIs on multivariable analysis. The majority of our DSC cohort had undergone
- arterial switch operations and hypoplastic aortic arch repairs with high RACHS-1 scores of 4,
- similar to several other series [6, 24, 25]. This may be explained as more complex operations
- 18 with higher RACHS-1 scores are more likely to lead to myocardial oedema, coagulopathy and
- 19 haemodynamic instability, increasing the risk of infection.
- 20 Application of ECMO was a significant risk factor for 30-day mortality in our multivariable
- analysis, similar to previous reports [26]. Gupta et al in a 998 paediatric cohort describe a 48.1%
- survival rate to hospital discharge with ECMO, with a mortality rate increasing by 12% every
- 23 day after 7 days of veno-arterial ECMO use[26].

Timing of sternal closure is still a matter of controversy. Riphagen et al. recommends DSC within 24 hours to lower rates of nosocomial infections and ventilator associated complications[24]. However, premature closure of the sternum can lead to repeated DSCs leading to more deleterious effects than a prolonged period of DSC[7]. Our study reports a DSC time of 3.8 (SD 5.8) days, comparable to the existing literature averaging at 3 days[17, 18]. Furthermore, our study reports no association between DSC time and risk of SWI. This is contradictory to previous retrospective studies which reported a higher incidence of infection with longer periods of open chest[2, 27]. This demonstrates the importance of avoiding

premature closure of the sternum as premature closure may lead to further attempts at

Limitations

reopening the chest[7, 18].

It is important to note that although several retrospective studies have been referenced, their outcomes cannot be generalised or used as a basis for comparison due to the significant variance in reported mortality and morbidity. Limitations of our study include it being retrospective and a single centre. Moreover, different surgical delayed sternal closure protocols exist within our surgeons' group.

# 1 Conclusion

- Delayed sternal closure (DSC) represents an efficacious surgical approach for the management
  of complex cardiac surgery in the paediatric population. Reassuringly, our findings reveal
- 4 minimal incidence of sternal wound infection associated with delayed sternal closure in these
- 5 high-risk cases. High RACHS-1 score was the only independent risk factor for sternal wound
- 6 infection in univariate analysis (p = 0.019) and multivariable analysis (p = 0.05). DSC facilitates
- 7 PAB adjustment in high-risk complex cases. Moreover, DSC allows quick access to initiate
- 8 postoperative lifesaving ECMO in this critical group of patients.

9

10

11

# **Funding**

No funding was received for this research.

13

14

# **Competing interests**

- All authors declare no conflict of interest. No authors have any personal or institutional
- 16 financial interests related to any content of this manuscript.

17

18

19

20

21

#### References

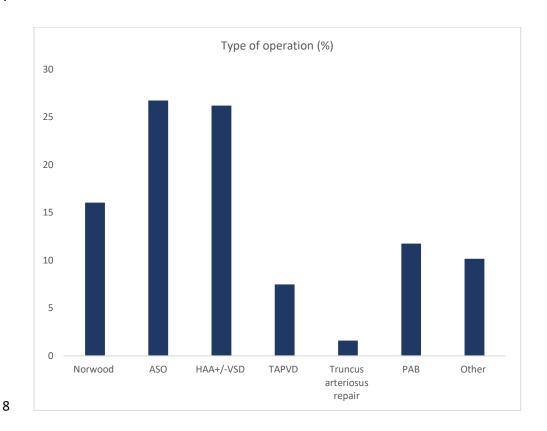
2	[1]	von Stumm M, Leps Y, Jochheim L, et al. Impact of delayed sternal closure on wound
3		infections following neonatal and infant cardiac surgery. PLoS One 2022; 17:
4		e0267985.

- Nelson-McMillan K, Hornik CP, He X, et al. Delayed Sternal Closure in Infant Heart Surgery-The Importance of Where and When: An Analysis of the STS Congenital Heart Surgery Database. *Ann Thorac Surg* 2016; 102: 1565–1572.
- [3] Hurtado-Sierra D, Calderón-Colmenero J, Curi-Curi P, et al. Outcomes of Delayed
  Sternal Closure in Pediatric Heart Surgery: Single-Center Experience. *Biomed Res Int* 2018; 2018: 3742362.
- 11 [4] Yabrodi M, Hermann JL, Brown JW, et al. Minimization of Surgical Site Infections in 12 Patients With Delayed Sternal Closure After Pediatric Cardiac Surgery. World J 13 Pediatr Congenit Heart Surg 2019; 10: 400–406.
- 14 [5] Gielchinsky I, Parsonnet V, Krishnan B, et al. Delayed sternal closure following open-15 heart operation. *Ann Thorac Surg* 1981; 32: 273–277.
- Özker E, Saritaş B, Vuran C, et al. Delayed sternal closure after pediatric cardiac
  operations; single center experience: a retrospective study. *J Cardiothorac Surg* 2012;
  7: 102.
- Harder EE, Gaies MG, Yu S, et al. Risk factors for surgical site infection in pediatric cardiac surgery patients undergoing delayed sternal closure. *J Thorac Cardiovasc Surg* 2013; 146: 326–333.
- Mehta PA, Cunningham CK, Colella CB, et al. Risk factors for sternal wound and other infections in pediatric cardiac surgery patients. *Pediatr Infect Dis J* 2000; 19: 1000–1004.
- Bowman ME, Rebeyka IM, Ross DB, et al. Risk factors for surgical site infection after delayed sternal closure. *Am J Infect Control* 2013; 41: 464–465.
- [10] Delgado-Corcoran C, Van Dorn CS, Pribble C, et al. Reducing Pediatric Sternal
  Wound Infections: A Quality Improvement Project. *Pediatr Crit Care Med* 2017; 18:
  461–468.
- 30 [11] Shin HJ, Jhang WK, Park J-J, et al. Impact of delayed sternal closure on postoperative infection or wound dehiscence in patients with congenital heart disease. *Ann Thorac* 32 Surg 2011; 92: 705–709.
- Adler AL, Smith J, Permut LC, et al. Significance of positive mediastinal cultures in pediatric cardiovascular surgical procedure patients undergoing delayed sternal closure. *Ann Thorac Surg* 2014; 98: 685–690.
- Kumar SR, Scott N, Wells WJ, et al. Liberal Use of Delayed Sternal Closure in
  Children Is Not Associated With Increased Morbidity. *Ann Thorac Surg* 2018; 106:
  581–586.

- 1 [14] Mills KI, van den Bosch SJ, Gauvreau K, et al. Physiologic effects of delayed sternal closure following stage 1 palliation. *Cardiol Young* 2018; 28: 1393–1403.
- [15] Woodward C, Taylor R, Son M, et al. Multicenter Quality Improvement Project to
  Prevent Sternal Wound Infections in Pediatric Cardiac Surgery Patients. World J
  Pediatr Congenit Heart Surg 2017; 8: 453–459.
- [16] Yang Y, Wang J, Cai L, et al. Surgical site infection after delayed sternal closure in neonates with congenital heart disease: retrospective case-control study. *Ital J Pediatr* 2021; 47: 182.
- 9 [17] Tabbutt S, Duncan BW, McLaughlin D, et al. Delayed sternal closure after cardiac operations in a pediatric population. *J Thorac Cardiovasc Surg* 1997; 113: 886–893.
- 11 [18] Elassal AA, Eldib OS, Dohain AM, et al. Delayed Sternal Closure in Congenital Heart 12 Surgery: A Risk-Benefit Analysis. Heart Surg Forum 2019; 22: E325–E330.
- 13 [19] Velho TR, Pereira RM, Guerra NC, et al. The impact of cardiopulmonary bypass time 14 on the Sequential Organ Failure Assessment score after cardiac surgery. Interdiscip 15 Cardiovasc Thorac Surg 2024; 38: ivae082.
- 16 [20] Biglioli P, Cannata A, Alamanni F, et al. Biological effects of off-pump vs. on-pump 17 coronary artery surgery: focus on inflammation, hemostasis and oxidative stress. Eur J 18 Cardiothorac Surg 2003; 24: 260–269.
- 19 [21] Kraft F, Schmidt C, Van Aken H, et al. Inflammatory response and extracorporeal circulation. Best Pract Res Clin Anaesthesiol 2015; 29: 113–123.
- 21 [22] Landis RC. Redefining the systemic inflammatory response. Semin Cardiothorac Vasc 22 Anesth 2009; 13: 87–94.
- [23] Ruggieri VG, Bounader K, Verhoye JP, et al. Prognostic Impact of Prolonged Cross Clamp Time in Coronary Artery Bypass Grafting. Heart Lung Circ 2018; 27: 1476–
  1482.
- 26 [24] Riphagen S, McDougall M, Tibby SM, et al. 'Early' delayed sternal closure following pediatric cardiac surgery. *Ann Thorac Surg* 2005; 80: 678–684.
- 28 [25] Samir K, Riberi A, Ghez O, et al. Delayed sternal closure: a life-saving measure in neonatal open heart surgery; could it be predictable? *Eur J Cardiothorac Surg* 2002; 21: 787–793.
- Gupta P, Robertson MJ, Beam B, et al. Relationship of ECMO duration with outcomes after pediatric cardiac surgery: a multi-institutional analysis. *Minerva Anestesiol* 2015; 81: 619–627.
- Anderson CA, Filsoufi F, Aklog L, et al. Liberal use of delayed sternal closure for postcardiotomy hemodynamic instability. *Ann Thorac Surg* 2002; 73: 1484–1488.

# 2 Figures:

**Figure 1:** Types of operations (%). ASO: Arterial Switch operation, HAA +/- VSD: Hypoplastic aortic arch repair with or without ventricular septal defect repair, TAPVD: Total anomalous pulmonary venous drainage repair, PAB: pulmonary artery banding



# 2 Tables:

# 3 Table 1: Demographic data

Variables	All cases (n = 187)
Age at operation in days (mean, SD)	$12.86 \pm 6.45$
Female gender (N, %)	59 (31.6%)
RACHS-1 score (mean, SD)	$4 \pm 1.05$
Weight at surgery in kg (mean, SD)	$3.33 \pm 0.48$
ICU stay in days (mean, SD)	$12.75 \pm 16.57$
Hospital stay in days (mean, SD)	$25.93 \pm 36.88$
Time to DSC in days (mean, SD)	$3.81 \pm 5.80$
Pre-operative ECMO (N, %)	1 (0.5%)
Operative ECMO (N, %)	6 (3.2%)
Postoperative ECMO (N, %)	21 (11.2%)
Cardiopulmonary bypass time in minutes (mean, SD)	$183.1 \pm 80.94$
Aortic cross clamp time in minutes (mean, SD)	$102.75 \pm 52.50$
30-day mortality (N, %)	9 (4.3 %)
Sepsis (N, %)	24 (12.8%)
Chest explorations (N, %)	52 (27.8%)

Table 2: Univariate logistic regression analysis for dependent variable, sternal wound infections. CI= confidence interval. The bold values show statistical significance (p <0.05).

Variables	p-value	95%	C.I.
Age at operation (days)	.139	006	.007
Sex	.168	156	.027
Weight at operation	.462	129	.059
ICU days	.987	003	.003
DSC days	.074	025	.001
ЕСМО	.299	248	.077
RACHS-1 (surgical risk score)	.019	.009	.101
Aortic cross clamp time	.388	003	.001
CPB time	.07	.000	.003
Number of chest explorations	.251	047	.178

**Table 3:** Univariate logistic regression analysis for dependent variable, 30- day mortality rate. CI= confidence interval. The bold values show statistical significance (p < 0.05).

Variables	p-value	95% C.I.	
Age at operation (days)	.583	005	.003
Sex	.833	067	.054
Operative weight	.845	068	.056
ICU days	.207	003	.001
DSC days	.011	019	003
ECMO	.000	.251	.465
RACHS-1 (surgical risk	.665	037	.024
score)			
Aortic cross clamp time	.007	.000	.003
CPB time	.006	002	.000
Number of chest	.215	027	.121
explorations			

**Table 4:** Multivariable logistic regression analysis for risk factors of sternal wound infections among neonates undergoing DSC post cardiac surgery. CI= confidence interval. The bold values show statistical significance (p < 0.05).

Variables	p-value	95%	C.I.
Age at operation (days)	.722	.925	1.120
Sex	.237	.607	7.533
Weight at operation	.49	.158	2.417
ICU days	.82	.964	1.048
DSC days	.211	.768	1.060
ECMO	.232	.450	26.953
RACHS-1 (surgical risk	.05	.995	4.012
score)			
Aortic cross clamp time	.536	.972	1.015
CPB time	.165	.995	1.030
Number of chest	.347	.526	6.210
explorations			

**Table 5:** Multivariable logistic regression analysis for risk factors of 30-day mortality among neonates undergoing DSC post cardiac surgery. CI= confidence interval. The bold values show statistical significance (p < 0.05).

Variables	p-value	95%	C.I.
Age at operation (days)	.359	.666	1.159
Sex	.336	.180	150.776
Operative weight	.629	.023	9.672
ICU days	.116	.680	1.043
DSC days	.824	.648	1.412
ECMO	.002	.000	.122
RACHS-1 (surgical	.126	.029	1.550
risk score)			
Aortic cross clamp	.159	.987	1.086
time			
CPB time	.604	.966	1.020
Number of chest	.915	.068	20.174
explorations			