



ELSEVIER

Available online at [www.sciencedirect.com](http://www.sciencedirect.com)

Journal of Hospital Infection

journal homepage: [www.elsevier.com/locate/jhin](http://www.elsevier.com/locate/jhin)



# Identifying potential predictors of the risk of surgical site infection following cardiac surgery: a scoping review

K.V. Charlwood<sup>a,\*</sup>, J. Jackson<sup>a</sup>, R. Vaja<sup>b</sup>, L.J. Rogers<sup>c</sup>, S. Dawson<sup>a</sup>,  
K.R. Moawad<sup>d</sup>, J. Brown<sup>e</sup>, J. Trevis<sup>f</sup>, I. Vokshi<sup>g</sup>, G.R. Layton<sup>h</sup>, R. Magboo<sup>i</sup>,  
J. Tanner<sup>j</sup>, M. Rochon<sup>k</sup>, G.J. Murphy<sup>l</sup>, P. Whiting<sup>a</sup>

<sup>a</sup> Population Health Sciences, Bristol Medical School, University of Bristol, Bristol, UK

<sup>b</sup> Royal Brompton Hospital, National Heart and Lung Institute, Imperial College London, London, UK

<sup>c</sup> Bristol Heart Institute, University Hospitals Bristol & Weston NHS Foundation Trust, Bristol, UK

<sup>d</sup> University Hospital Southampton Trust, Southampton, UK

<sup>e</sup> Royal Victoria Hospital, Belfast, UK

<sup>f</sup> Freeman Hospital, Newcastle-upon-Tyne, UK

<sup>g</sup> Heart and Lung Centre, New Cross Hospital, Wolverhampton, UK

<sup>h</sup> University of Leicester, Leicester, UK

<sup>i</sup> Barts Heart Centre, St Bartholomew's Hospital, Barts Health NHS Trust, London, UK

<sup>j</sup> School of Health Sciences, University of Nottingham, Nottingham, UK

<sup>k</sup> Guy's and St Thomas NHS Foundation Trust, London, UK

<sup>l</sup> Leicester NIHR Biomedical Research Centre, University of Leicester, Leicester, UK

## ARTICLE INFO

### Article history:

Received 30 August 2024

Received in revised form  
30 November 2024

Accepted 4 December 2024

Available online 15 December  
2024

### Keywords:

Cardiac surgery  
Surgical site infection  
Risk prediction  
Scoping review

## SUMMARY

**Objectives:** This scoping review was undertaken to identify risk prediction models and pre-operative predictors of surgical site infection (SSI) in adult cardiac surgery. A particular focus was on the identification of novel predictors that could underpin the future development of a risk prediction model to identify individuals at high risk of SSI, and therefore guide a national SSI prevention strategy.

**Methods:** A scoping review to systematically identify and map out existing research evidence on pre-operative predictors of SSI was conducted in two stages. Stage 1 reviewed prediction modelling studies of SSI in cardiac surgery. Stage 2 identified primary studies and systematic reviews of novel cardiac SSI predictors.

**Results:** The search identified 7887 unique reports; 7154 were excluded at abstract screening and 733 were selected for full-text assessment. Twenty-nine studies (across 30 reports) were included in Stage 1 and reported the development ( $N=14$ ), validation ( $N=13$ ), or both development and validation ( $N=2$ ) of 52 SSI risk prediction models including 67 different pre-operative predictors. The remaining 703 reports were re-assessed in Stage 2; 49 studies met the inclusion criteria, and 56 novel pre-operative predictors that have not been assessed previously in models were identified.

This article is part of a special issue entitled: WHO Issue: IPC and AMS published in Journal of Hospital Infection.

\* Corresponding author. Address: NIHR Applied Research Collaboration West (NIHR ARC West), 9th Floor, Whitefriars, Lewins Mead, Bristol BS1 2NT, UK.

E-mail address: [katie.charlwood@bristol.ac.uk](mailto:katie.charlwood@bristol.ac.uk) (K.V. Charlwood).

<https://doi.org/10.1016/j.jhin.2024.12.002>

0195-6701/© 2024 The Author(s). Published by Elsevier Ltd on behalf of The Healthcare Infection Society. This is an open access article under the CC BY license (<http://creativecommons.org/licenses/by/4.0/>).



**Conclusions:** This review identified 123 pre-operative predictors of the risk of SSI following cardiac surgery, 56 of which have not been included previously in the development of cardiac SSI risk prediction models. These candidate predictors will be a valuable resource in the future development of risk prediction scores, and may be relevant to prediction of the risk of SSI in other surgical specialities.

© 2024 The Author(s). Published by Elsevier Ltd on behalf of The Healthcare Infection Society. This is an open access article under the CC BY license (<http://creativecommons.org/licenses/by/4.0/>).

## Introduction

Surgical site infections (SSIs) are the most common healthcare-associated infections in people undergoing cardiac surgery in the UK [1]. They are estimated to affect approximately 8.6% of patients undergoing coronary artery bypass graft (CABG) surgery and 2.2% of patients undergoing non-CABG surgery up to 30 days post-operatively [2]. SSIs are associated with a 10-fold increase in mortality, a six-fold increase in hospital re-admission, prolonged hospitalisation [3], the need for further surgery, and extended outpatient care [4]. The direct costs of treating SSIs are estimated to exceed £15 million annually in the UK [5]. Targeted SSI prevention strategies may be as clinically effective as non-targeted interventions, more cost-effective, and reduce the drivers of antimicrobial resistance. To date, no randomised controlled trials testing such interventions exist.

Risk prediction modelling can be utilised to predict future outcomes following surgery. Pre-operative identification of patients with a high risk of SSI can guide the use of prevention and treatment strategies to reduce this risk, or prompt early intervention to prevent disease progression. These prediction models can consider a range of factors that are often patient related, such as age, comorbidities and the surgical intervention itself. Most models for cardiac surgery populations have been developed to predict mortality and general morbidity outcomes [6,7], rather than SSI specifically. Furthermore, there are limitations to how these models have been validated, with most only being validated in a specific geographic or surgical population. The performance of even those models with the greatest evidence base in predicting SSI in cardiac surgery is limited; the Australian Clinical Risk Index [8] and Brompton Harefield Infection Score [9] have area under the curve (AUC) values of approximately 0.7. Currently, the use of risk prediction models to guide SSI prevention strategies remains inconsistent [2].

This scoping review was undertaken to identify candidate pre-operative predictors of SSI in adults following cardiac surgery that have been considered during the development of existing risk prediction models, and novel predictors from primary studies that have not been included in any previous prediction models. These candidate predictors will be considered in the development of a future risk prediction model of SSI following cardiac surgery that will underpin the identification of high-risk patients. In combination with work to identify barriers and facilitators to SSI prevention and surveillance [10,11], a national SSI prevention strategy in adult cardiac surgery will be developed.

## Methods

This scoping review was conducted to systematically identify and map out pre-operative predictors of SSI following

cardiac surgery. The review was conducted in two stages, using the same search for both. Stage 1 identified prediction modelling studies of SSI in patients who underwent cardiac surgery, and Stage 2 identified primary studies and systematic reviews of novel predictors of SSI in these patients. This scoping review has been reported according to the PRISMA Extension for Scoping Reviews [12]. The original protocol was registered on the PROSPERO database [13]; protocol amendments are described in [Appendix A](#) (see online supplementary material).

### Sources of evidence

Embase, MEDLINE and Web of Science were searched from 2000 to February 2022, June 2022 and July 2022, respectively. There have been substantial changes in the use of antimicrobials in clinical practice over the past few decades, so the searches were restricted to reports since 2000 for applicability. The database searches were carried out sequentially, allowing for each subsequent search to be recalibrated to optimise sensitivity and specificity. The search strategy combined terms for cardiac surgery and SSI together with a bespoke filter to identify candidate risk predictors or predictive models ([Appendix B](#), see online supplementary material). The search also included a parallel search strand for known predictors of SSI or existing cardiac surgery risk scores. The search results were limited to studies in adults by applying a filter to remove non-human studies, and research in neonates, infants or children. Case reports, editorials and letters were excluded, and no language restrictions were applied. The reference lists of relevant guidelines, systematic reviews and included studies were also screened to identify any additional reports.

### Stage 1. Identifying pre-operative predictors in risk prediction modelling studies

#### Study selection

Prediction modelling studies on risk of SSI in adults (age  $\geq 18$  years) undergoing clean operations in cardiac surgery which assessed at least one pre-operative predictor were eligible. A prediction modelling study was defined based on the PROBAST definition [14] as 'a study that aimed to develop, validate, or update a multi-variable prediction model to estimate the risk of the occurrence of SSI in adults following clean cardiac surgery'.

The population of interest included patients undergoing clean cardiac surgery. Clean surgery was defined using the Centers for Disease Control and Prevention classification of Class 1 wounds: uninfected, no inflammation present, and closed primarily. If wound drainage was required, a closed draining method was necessary [15]. Any cardiac surgeries via median sternotomy were eligible, but surgeries for infective

endocarditis, an infective source (abscess, wound, graft infection), and minimally invasive surgeries were excluded.

SSI was defined as any infection originating in cardiac surgical wounds or the organs or spaces opened or manipulated during the operative procedure [16]. Studies that reported this outcome as (but not limited to) any SSI, mediastinitis or deep sternal wound infection (DSWI) were included. Studies that reported SSI only as part of a composite outcome were excluded.

Search results were screened independently by two reviewers to identify any potentially eligible reports for either review stage. Studies considered relevant for Stage 1 were obtained for full-text review, and assessed for inclusion in Stage 1 independently by two reviewers. Disagreements were resolved through discussion or referral to a third reviewer.

#### Data extraction

Data were extracted into standardized forms developed in Excel (Microsoft Corp, Redmond, WA, USA). These forms were piloted on a small number of studies and adapted as necessary. Data were extracted on study characteristics, patient characteristics, type of surgery, SSI definition, and follow-up period. All patient-level predictors that could be measured

prior to surgery were extracted and mapped. Predictors that could not be measured prior to surgery (e.g. duration of operation, length of ICU stay) or were related to surgical preparation (e.g. method of hair removal, bacterial decolonisation) were excluded as these may be influenced by facility-specific procedures.

The type of prediction modelling study, the modelling methods used, what eligible predictors were assessed and included in the final model, and the discriminative capability of the model (AUC) were also extracted. For studies that included multiple prediction models, data were extracted for all models that met the review eligibility criteria. Models with the same name included in multiple studies that were validated in a different surgical population, or used a different outcome definition, were classified as distinct models, and data were extracted separately for these.

#### Synthesis

The results were described and charted based on existing scoping frameworks and guidance [17,18]. The key study characteristics and prediction model characteristics were depicted using tables and charts. Simplifications were made to aid predictor mapping and understanding of their usage in

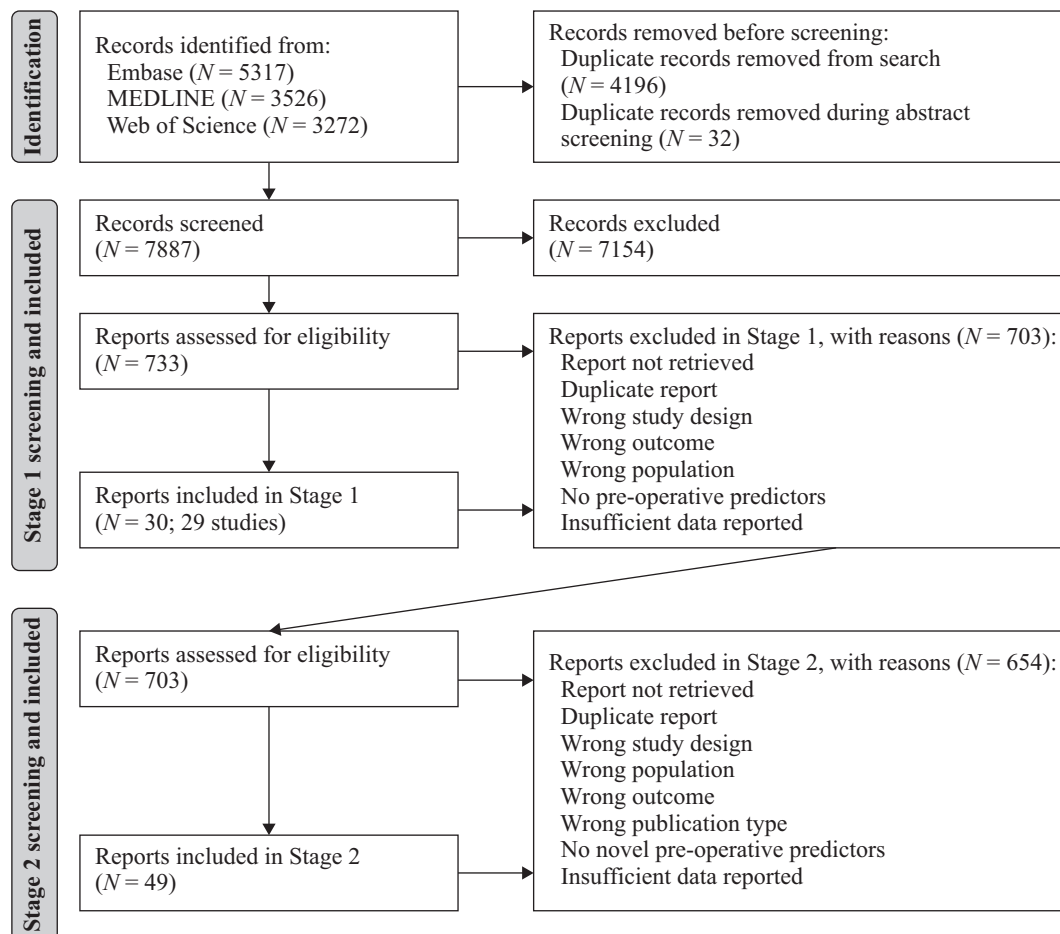


Figure 1. PRISMA flow diagram displaying the two-stage screening approach.

existing cardiac SSI risk prediction models. Predictors with similar or overlapping definitions were grouped together; for example, 'mitral insufficiency' and 'mitral stenosis' were grouped as 'mitral valve disease'. All extracted pre-operative predictors were presented in a table, along with the frequencies of assessment and inclusion across all included models. To understand how the broader clinical categories of predictors have been used in these existing models, individual predictors were organised into four categories: medication, biochemistry, demographics and comorbidity. The discriminative performance (AUC) of models across different studies has also been tabulated.

## Stage 2. Identifying novel pre-operative predictors in primary studies

### Study selection

Stage 2 was conducted after data extraction was completed for Stage 1. Primary studies of a cohort or case–control design and systematic reviews that evaluated the association between a 'novel' predictor and SSI in adults (age  $\geq 18$  years) undergoing clean cardiac surgery were included. Cohort studies measured one or more predictors in a cohort of patients undergoing cardiac surgery who were then followed-up to determine whether or not they developed SSI. Case–control studies selected patients with SSI following cardiac surgery and patients without SSI following cardiac surgery, comparing predictor frequencies between groups. Novel predictors were defined as any factor that could be measured pre-operatively that had not been evaluated in any of the prediction modelling studies included in Stage 1.

To identify relevant studies for Stage 2, two reviewers independently rescreened the full texts of all studies that were excluded from Stage 1. The two reviewers assessed and included studies that evaluated eligible predictors.

### Data extraction

As with Stage 1, data were extracted into standardised, piloted forms developed in Excel. Data on study characteristics, patient characteristics, type of surgery outcome definition, and length of follow-up were extracted. 2x2 tables were constructed showing the number of participants with and without SSI, cross-classified against the number with and without the novel candidate predictor.

### Synthesis

The key characteristics from the extracted studies were tabulated. The extracted novel predictors were listed in a table, along with *P*-values, to indicate evidence of an association with the risk of SSI following cardiac surgery.

## Results

### Search results

The searches identified 7887 unique records; 7154 studies were excluded during title and abstract screening, and 733 studies were selected for full-text assessment. Twenty-nine studies (detailed in 30 reports) were included in Stage 1. The remaining 703 studies were re-assessed for inclusion in Stage 2;

49 studies were included in Stage 2. [Figure 1](#) summarizes the screening approach.

## Stage 1. Pre-operative predictors included in risk prediction modelling studies

### Study characteristics

Twenty-nine studies reported the development ( $N=14$ ), validation ( $N=13$ ), or development and validation ( $N=2$ ) of 52 prediction models of the risk of SSI following cardiac surgery. [Table 1](#) outlines the key study characteristics. Most studies were conducted in European or North American countries (72%). Almost all were cohort studies (93%) that assessed at least 1000 patients (86%). For studies that reported the duration of follow-up, most followed-up for 30 days after surgery (20%).

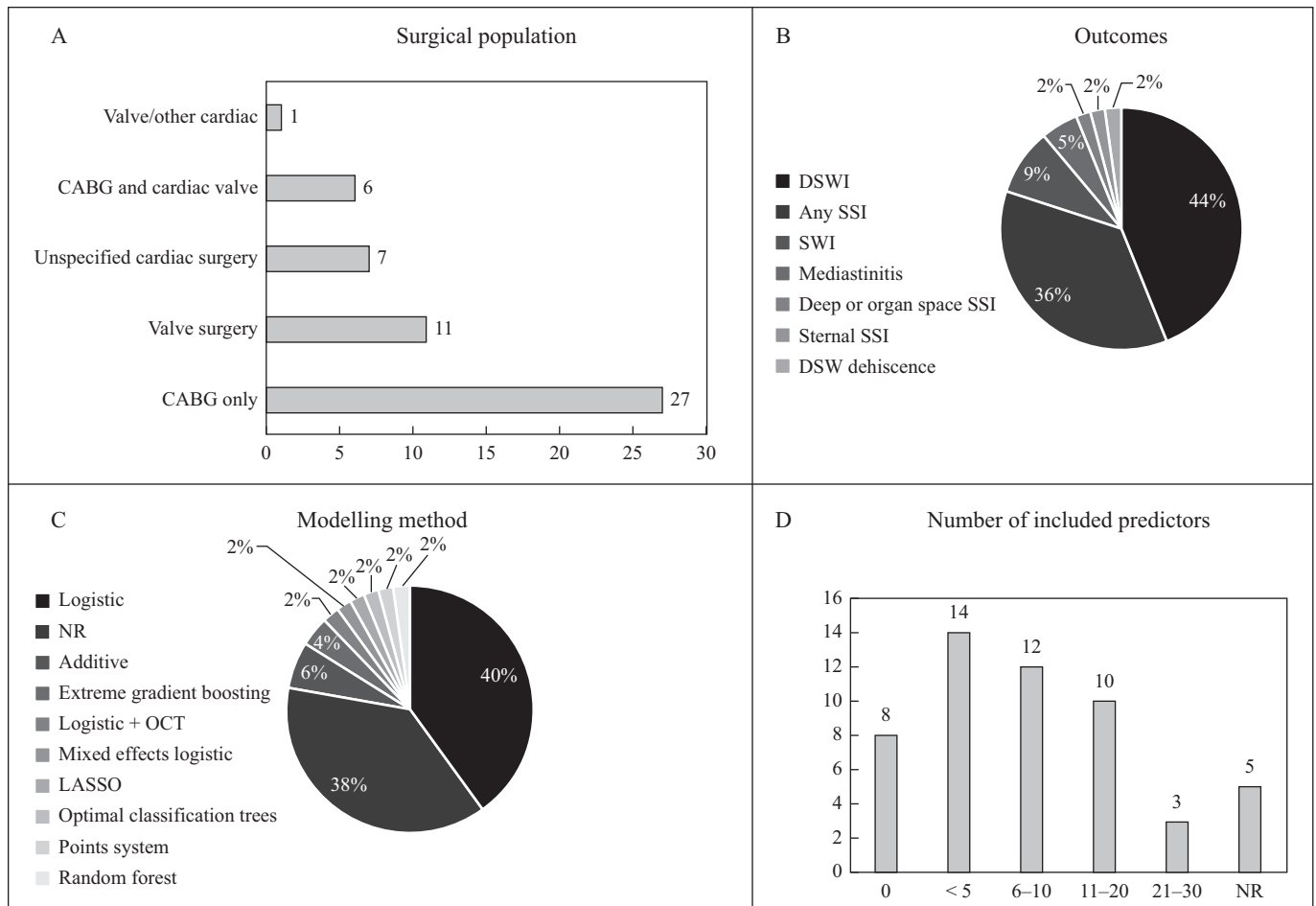
### Model characteristics

Of the 52 SSI risk prediction models, 40 included pre-operative predictors alone, and 12 also included intra-operative predictors. [Figure 2](#) describes the key characteristics of these models. Twenty-four models (46%) predicted the risk of SSI in patients who had undergone isolated CABG surgery, with other models commonly developed for isolated valve surgery (21%) and mixed cardiac surgery (14%).

There was considerable variation in the SSI definitions used ([Figure 2b](#)). Most models were for DSWI (44%) or any SSI (36%) outcomes. Logistic regression was most commonly used in model development (40%). Models developed more recently used a greater diversity of methods, including machine learning approaches (4% used extreme gradient boosting).

**Table 1**  
Characteristics of included prediction modelling studies

Characteristic	Category	Number of studies
Study type	Development	14
	Validation	13
	Development and validation	2
Study design	Cohort	27
	Nested case–control	1
	Case–control	1
Location	North America	12
	Europe	9
	Asia	3
	Australia and Oceania	3
	South America	1
Follow-up time (days)	International	1
	30	7
	>30	1
	60	1
	Until discharge	2
	Postoperative period	1
	NR	17
Sample size	100–999	8
	1000–9999	15
	10,000–99,999	3
	100,000–999,999	7



**Figure 2.** Overview of model characteristics. Figure panel includes surgical population (A), outcome of surgical site infection (SSI) (B), prediction modelling method used (C), and number of pre-operative factors included in the final model (D). NR, not recorded (if the characteristic could not be identified in the study report). CABG, coronary artery bypass graft; DSWI, deep sternal wound infection; SWI, sternal wound infection; OCT, optimal classification trees.

Approximately half of the final models included more than 10 pre-operative predictors, with the total number ranging from one to 26. Eight finalised models included intra-operative predictors only. It was not possible to identify the predictors included in the models ( $N=5$ ) developed by Orfanoudaki *et al.* as these were not reported in the development study [19].

#### Pre-operative predictors

Table II summarises the pre-operative predictors considered for inclusion, and those selected for the final prediction models. The most common predictors included in the final models were comorbidities and demographic factors. These included body mass (69%), diabetes (50%), gender (38%), chronic obstructive pulmonary disease (38%), age (35%), peripheral vascular disease (33%), cardiogenic shock (29%), left ventricular ejection fraction (27%), surgical priority (27%), and number of diseased vessels (23%). The pre-operative predictors included in the finalised model are reported in Appendix C, Table III (see online supplementary material).

Most models were found to have low-to-moderate discriminative performance [ $AUC=0.5-0.7$ , Appendix C, Table IV (see

online supplementary material)]. The STS model had the greatest reported performance ( $AUC=0.89$ ) in a study validating a prediction model for the risk of DSWI in an Indian cardiac surgery population [20].

#### Stage 2. Novel Pre-operative predictors included in primary studies

##### Study characteristics

Table III summarises the characteristics of the studies included in Stage 2. Most studies were conducted in European or North American countries (65%). Thirty-four (69%) were cohort studies, 13 (27%) were case–control studies, and two were systematic reviews. Cardiac surgery (49%) and isolated CABG surgery (43%) were the most common study populations. Outcome definition varied and included DSWI, mediastinitis, all SSIs, sternal wound infection and wound infection. Follow-up ranged from 30 days to 3 years following surgery; the majority of studies followed-up for 30 days. Patient sample sizes ranged from 23 to 5.6 million, with studies most commonly assessing 100–1000 patients (49%).

**Table II**

Pre-operative predictors in existing models of risk of surgical site infection following cardiac surgery

Predictor	Included	Assessed
Biochemistry		
Creatinine	11	9
Haematocrit	1	6
Platelet count	1	6
White blood cell count	1	1
Haemoglobin	0	5
Albumin	0	1
Comorbidities		
Diabetes	26	5
Chronic obstructive pulmonary disease	20	7
Peripheral vascular disease	17	5
Cardiogenic shock	15	10
Surgical priority	14	14
Left ventricular ejection fraction	14	11
Number of diseased vessels	12	7
Renal impairment	10	13
Heart failure	10	9
Endocarditis	10	9
Previous cardiac surgery	9	15
Myocardial infarction	9	12
Angina	9	10
Cerebrovascular event	7	13
Hypertension	6	9
Pulmonary hypertension	6	7
Type of surgery	6	4
Neurological disorder	5	7
ASA score	4	5
Arrhythmia	3	8
Valvular disease (mitral)	2	9
Valvular disease (aortic)	1	10
Valvular disease (tricuspid)	1	9
Valvular disease (pulmonic)	1	5
Intubation	1	2
Left ventricular size	1	0
Pulmonary function	1	0
Malignancy	0	7
Chest infection	0	7
Thoracic radiotherapy	0	7
Hepatic disease	0	7
Syncope	0	7
Left ventricular dimension	0	5
Previous cardiac surgery (valve disease)	0	5
Colostomy	0	3
Coagulopathy	0	2
EuroSCORE II	0	2
Dependency	0	2
Weight loss	0	2
Obstructive sleep apnoea	0	2
Hypercholesterolaemia	0	1
Anaemia	0	1
Deep venous thrombosis	0	1
Demographics		
Body mass	36	5
Gender	20	10
Age	18	9
Surgical period	8	8
Ethnicity	6	6
Smoking	5	11

**Table II (continued)**

Predictor	Included	Assessed
Alcohol consumption	1	6
Family history of heart disease	1	6
Socio-economic status	1	1
Recreational drug use	0	7
Pre-operative length of stay	0	5
Medications		
Immunosuppressants	5	10
ACE inhibitors	0	7
ADP receptor inhibitors	0	7
Glycoprotein IIb/IIIa inhibitors	0	7
Anticoagulants	0	5
Antiplatelets	0	5
Beta blockers	0	5

ASA, American Society of Anesthesiologists; ACE, angiotensin-converting enzyme; ADP, adenosine diphosphate.

Assessed, assessed but not included in model; included, assessed and included in model (not counted in 'assessed'). If two or more similar predictors were assessed or included in the same model, this was counted once.

**Table III**

Key characteristics of studies included in Stage 2

Characteristic	Category	Number of studies
Study design	Cohort	34
	Case-control	13
	Systematic review	2
Location	Europe	18
	North America	14
	Asia	9
	South America	4
	Australia/Oceania	2
	International	2
Surgical population	Cardiac	24
	CABG	21
	CABG and valve	1
	CABG and/or valve	2
	CABG or SAVR	1
Outcome	DSWI	14
	Mediastinitis	12
	SSI	12
	SWI	11
	Wound infection	3
	Follow-up period	30 days
>30 days	2	
90 days	4	
6 months	1	
1 year	1	
3 years	1	
NR	31	
Sample size	<100	7
	100–999	24
	1000–9999	11
	10,000–99,999	3
	100,000–999,999	2
	≥1 million	2

CABG, coronary artery bypass graft; SAVR, surgical aortic valve replacement; DSWI, deep sternal wound infection; SSI, surgical site infection; SWI, sternal wound infection; NR, not reported.

Total outcome count is 52 as three studies examined two SSI outcomes separately.

Table IV

Novel pre-operative factors and their association with risk of surgical site infection (SSI) following cardiac surgery: supporting and contradictory studies

Predictor	Associated with risk of SSI	No association
Biochemistry		
Bilirubin	–	Theodore, 2019 [41]
Blood urea	Bugra, 2021 [26]	–
Calcium	–	Bugra, 2021 [26]
Cholesterol	–	Bugra, 2021 [26]
CRP	Bugra, 2021 [26]; Cappabianca, 2006 [27]; Elenbaas, 2010 [28]	Tschudin–Sutter, 2013 [29]
ESR	Bugra, 2021 [26]; Togan, 2015 [42]	–
HDL	Bugra, 2021 [26]	–
LDL	–	Bugra, 2021 [26];
GFR	–	Theodore, 2019 [41]
Iron deficiency	–	Immohr, 2021 [43]
Oxidative stress	–	Suehiro, 2014 [44]
Comorbidities		
Asymptomatic bacteriuria	–	Duarte, 2018 [45]
Calcified aorta	Filsofi, 2009 [46]; Toumpoulis, 2005 [47]	–
Depression	Theodore, 2019 [41]	–
Frailty	Lemus-Barrios, 2020 [48]; Back, 2019 [49]	–
Gastrointestinal disease	–	Crape, 2021 [50]
Opiate addiction	Hosseinzadei, 2012 [51]	–
HIV	–	Boccaro, 2008 [34]; Dominici, 2020 [35]; Jimenez-Exposito, 2006 [36]; Robich, 2014 [37]
Hypothyroidism	–	Jaimés, 2017 [52]
Intramuscular adipose tissue content	Kiriya, 2020 [53]	–
Metabolic syndrome	Ozkan, 2017 [21]; Ozyazicioglu, 2010 [22]; Zapata, 2020 [23]	Ardeshiri, 2014 [24]; Pimenta, 2007 [25]
Pre-operative infection	–	Tadros, 2013 [54]; Zapata, 2020 [23]
Psoas total muscle index	–	Kiriya, 2020 [53]
Psychiatric history	–	Hassan, 2006 [55]
Rheumatoid arthritis	–	Hassan, 2006 [55]
<i>Staphylococcus aureus</i> colonization	Maillet, 2011 [56]; Munoz, 2008 [57]	–
MRSA colonization	Munoz, 2008 [57]	Cutrell, 2016 [58]; Dodds Ashley, 2004 [59]
UTI	–	Duarte, 2018 [45]
Vitamin B12	Bugra, 2021 [26]	–
Demographics		
Distressed Communities Index	Mehaffey, 2020 [60]	–
Hospital transfer	Al Salmi, 2019 [61]	–
Latitude	Abdelnoor, 2016 [62]	–
Payer status	–	Benedetto, 2021 [63]
Physical activity	Van Laar, 2017 (ages 66–75 years) [64]	Van Laar, 2017 (ages ≤ 65 and >75 years) [64]
Medication		
Alpha blockers	Eton, 2016 [30]	–
Aspirin	–	Eton, 2016 [30]; Robinson, 2007 [65]
Bronchodilators	–	Eton, 2016 [30]
Calcium channel blockers	–	Eton, 2016 [30]
Clopidogrel	–	Eton, 2016 [30]
Diuretics	–	Eton, 2016 [30]
Heparin	–	Cayci, 2008 [66]
Heparin or nitrates	Nespor, 2015 [67]	–
Nitrates	–	Eton, 2016 [30]; Toumpoulis, 2005 [47]
NSAIDs	–	Eton, 2016 [30]
Proton pump inhibitors	–	Eton, 2016 [30]

(continued on next page)

Table IV (continued)

Predictor	Associated with risk of SSI	No association
SSRI/SNRIs	–	Tully, 2012 [68]
Statins	Kayani, 2013 [31]	Eton, 2016 [30]; Oddsson, 2012 (studied in mediastinitis and SWI separately) [32]; Young, 2010 [33]
Thrombolysis	–	Toumpoulis, 2005 [47]
Vasopressors	–	Eton, 2016 [30]
	Risk scores	
ACDS (continuous)	Batista, 2006 [69]	–
ACDS (ordinal, quintiles)	–	Batista, 2006 [69]
CHA <sub>2</sub> DS <sub>2</sub> -VASc score	Kalyoncuoglu, 2019 [70]	–
NHSN risk score (>1)	–	Cutrell, 2016 [58]
PACDS (continuous)	Batista, 2006 [69]	–
PACDS (ordinal, quintiles)	Batista, 2006 (Q3, Q5) [69]	Batista, 2006 (Q2, Q4) [69]
Prognostic nutritional index	Hayashi, 2020 [71]	Lee, 2020 [72]

ACDS, admission chronic disease score; CRP, C-reactive protein; GFR, glomerular filtration rate; HDL, high-density lipoprotein; HIV, human immunodeficiency virus; LDL, low-density lipoprotein; NHSN, National Healthcare Safety Network (USA); NSAID, non-steroidal anti-inflammatory drugs; PACDS, pre-admission chronic disease score; *S. aureus*, *Staphylococcus aureus*; MRSA, methicillin-resistant *Staphylococcus aureus*; ESR, erythrocyte sedimentation rate; SSRI, selective serotonin re-uptake inhibitors; SNRI, serotonin–norepinephrine re-uptake inhibitors; SWI, sternal wound infection; UTI, urinary tract infection.

Studies that found an association reported a *P*-value  $\leq 0.05$ .

### Novel pre-operative factors

Fifty-six pre-operative novel candidate predictors of the risk of SSI following cardiac surgery were identified (Table IV); most were comorbidities or related to medication use. The most commonly evaluated predictors in the included studies were metabolic syndrome [21–25], C-reactive protein (CRP) [26–29], use of statins [30–33], and human immunodeficiency virus (HIV) [34–37]. Findings on whether these predictors are associated with the risk of SSI were inconsistent, except for HIV where no association was found across all four studies (Table IV).

### Discussion

This review identified 52 existing prediction models of the risk of SSI following cardiac surgery that considered pre-operative predictors. Forty models included pre-operative predictors alone. Overall, model performance was found to be poor to moderate. Sixty-seven pre-operative predictors were considered during the development of these models, with final models most often including comorbidities and demographic predictors such as body mass and diabetes. Fifty-six novel pre-operative factors that have not been considered previously in cardiac SSI risk model development were identified. This review also highlights significant heterogeneity in the surgical populations sampled, duration of follow-up, and definitions of predictors and SSI outcomes across studies.

Over 100 predictors have been identified, but their correlations and interactions need to be assessed in a comparable population to evaluate their predictive utility. Given that a risk model with such a large number of variables is not feasible, a more pragmatic approach would involve using a large-scale national dataset to create a hierarchy of risk, identifying the top 10–20 variables, and developing a standardised scoring system.

Blood biomarkers have been included in cardiac SSI risk model development, but are often excluded from final models. This review identified several blood biomarkers of inflammation and lipid regulation that may improve prediction of the risk of SSI that have not been considered in model development. PhenoAge utilizes blood biomarkers such as CRP to estimate biological age [38], and has the potential to outperform the use of traditional risk factors in predicting morbidity and mortality. In the UK, blood testing is recommended prior to elective cardiac surgery [39]; the use of blood biomarkers in identifying the risk of SSI should be explored further.

The increase in antimicrobial-resistant infections requires careful decisions around the strategies implemented to mitigate risk. Given the increasing accessibility and wider use of electronic health records in research, future cardiac SSI risk prediction models should utilise routine data to predict not only SSI, but also organism type. A recent study in the *Journal of Hospital Infection* found that women are primarily susceptible to Gram-negative SSIs following cardiac surgery, whereas men are primarily susceptible to Gram-positive infections [40]. This has directly influenced antibiotic prophylaxis regimens within the study hospitals. Risk stratification tools would facilitate personalised SSI prevention strategies, potentially making them more cost-effective and reducing the burden of antimicrobial resistance.

As this was a scoping review, the aim was to identify risk prediction models and pre-operative predictors of SSI in adult cardiac surgery. The intention was not to provide a synthesis of the utility or accuracy of identified predictors. Risk of bias was not assessed, consistent with scoping review methodology [17,18]. Instead, this review focused on mapping the extent to which each predictor has been studied or considered in model development. Another limitation is the focus on cardiac surgery populations, potentially excluding predictors of the risk of SSI in other surgical populations that could be applicable to cardiac surgery. Furthermore, only patient-level predictors were included, as these are essential for individual risk



assessment. While pre-operative care practices, such as anti-microbial prophylaxis, affect patient risk, they are generally standardised across patients within a single setting, and are less suitable for individual risk stratification. The main strength of this review was the breadth of studies included, which facilitated the identification of pre-operative factors that have not been considered previously in risk modelling studies.

In conclusion, this review identified 123 pre-operative factors that could predict SSI following cardiac surgery, 56 of which have not been considered in existing risk prediction models. This list of candidate predictors will be valuable in the future development of risk prediction scores, and may also be relevant to other surgical specialities.

## Acknowledgements

The authors wish to thank the Cardiothoracic Interdisciplinary Research Network for their support with this review.

### Conflict of interest statement

None declared.

### Funding sources

This study was funded by a National Institute for Health Care Research (NIHR) programme development grant (NIHR202620). The views expressed are those of the authors and not necessarily those of NIHR.

### Ethical approval

Not required. This scoping review synthesizes publicly available information and does not involve patient data.

## Appendix A. Supplementary data

Supplementary data to this article can be found online at <https://doi.org/10.1016/j.jhin.2024.12.002>.

## References

- [1] Lamagni TCK, Wloch C, Harrington P. The epidemiology of cardiac surgical site infection in England, 2018/19. 30th European Congress of Clinical Microbiology and Infectious Diseases, Paris, 2020.
- [2] Cardiothoracic Interdisciplinary Research Network. National survey of variations in practice in the prevention of surgical site infections in adult cardiac surgery, United Kingdom and Republic of Ireland. *J Hosp Infect* 2020;106:812–9.
- [3] Gelijns AC, Moskowitz AJ, Acker MA, Argenziano M, Geller NL, Puskas JD, et al. Management practices and major infections after cardiac surgery. *J Am Coll Cardiol* 2014;64:372–81.
- [4] Joshi V, Vaja R, Richens D. Cost analysis of gentamicin-impregnated collagen sponges in preventing sternal wound infection post cardiac surgery. *J Wound Care* 2016;25:22–5.
- [5] Rochon M, Makecha S, Morais C, Luff D, Richardson Persaud-Rai B, Tibbles S, et al. A quality improvement approach to reducing readmission for surgical site infections. *Wounds UK* 2016;12:26–31.
- [6] Toumpoulis IK, Anagnostopoulos CE, Swistel DG, Derose Jr JJ. Does EuroSCORE predict length of stay and specific postoperative complications after cardiac surgery? *Eur J Cardiothorac Surg* 2005;27:128–33.
- [7] Shroyer ALW, Coombs LP, Peterson ED, Eiken MC, DeLong ER, Chen A, et al. The Society of Thoracic Surgeons: 30-day operative mortality and morbidity risk models. *Ann Thorac Surg* 2003;75:1856–65.
- [8] Figuerola-Tejerina A, Bustamante E, Tamayo E, Mestres CA, Bustamante-Munguira J. Ability to predict the development of surgical site infection in cardiac surgery using the Australian Clinical Risk Index versus the National Nosocomial Infections Surveillance-derived Risk Index. *Eur J Clin Microbiol Infect Dis* 2017;36:1041–6.
- [9] Rochon M, Jarman JW, Gabriel J, Butcher L, Morais C, Still M, et al. Multi-centre prospective internal and external evaluation of the Brompton Harefield Infection Score (BHIS). *J Infect Prev* 2018;19:74–9.
- [10] Tanner J, Brierley Jones L, Rochon M, Westwood N, Wloch C, Vaja R, et al. Barriers and facilitators for surgical site infection surveillance for adult cardiac surgery in a high-income setting: an in-depth exploration. *J Hosp Infect* 2023;141:112–8.
- [11] Tanner J, Brierley Jones L, Westwood N, Rochon M, Wloch C, Vaja R, et al. A comprehensive qualitative investigation of the factors that affect surgical site infection prevention in cardiac surgery in England using observations and interviews. *J Hosp Infect* 2024;149:119–25.
- [12] Tricco AC, Lillie E, Zarin W, O'Brien KK, Colquhoun H, Levac D, et al. PRISMA Extension for Scoping Reviews (PRISMA-ScR): checklist and explanation. *Ann Intern Med* 2018;169:467–73.
- [13] Rogers LJ, Vaja J, Jackson J, Dawson S, Whiting PF. Accuracy of candidate predictors and existing prediction models for surgical site infection (SSI) in clean surgery: a systematic review protocol. PROSPERO 2022 CRD42022309357. Available at: [https://www.crd.york.ac.uk/prospero/display\\_record.php?ID=CRD42022309357](https://www.crd.york.ac.uk/prospero/display_record.php?ID=CRD42022309357).
- [14] Wolff RF, Moons KGM, Riley RD, Whiting PF, Westwood M, Collins GS, et al. PROBAST: a tool to assess the risk of bias and applicability of prediction model studies. *Ann Intern Med* 2019;170:51–8.
- [15] Mangram AJ, Horan TC, Pearson ML, Silver LC, Jarvis WR. Guideline for prevention of surgical site infection, 1999. *Am J Infect Control* 1999;27:97–134.
- [16] Berríos-Torres SI, Umscheid CA, Bratzler DW, Leas B, Stone EC, Kelz RR, et al. Centers for Disease Control and Prevention guideline for the prevention of surgical site infection, 2017. *JAMA Surg* 2017;152:784–91.
- [17] Arksey H, O'Malley L. Scoping studies: towards a methodological framework. *Int J Soc Res Methodol Theory Pract* 2005;8:19–32.
- [18] Levac D, Colquhoun H, O'Brien KK. Scoping studies: advancing the methodology. *Implement Sci* 2010;5:69.
- [19] Orfanoudaki A, Giannoutsou A, Hashim S, Bertsimas D, Hagberg RC. Machine learning models for mitral valve replacement: a comparative analysis with the Society of Thoracic Surgeons risk score. *J Card Surg* 2022;37:18–28.
- [20] Borde D, Gandhe U, Hargave N, Pandey K, Khullar V. The application of European System for Cardiac Operative Risk Evaluation II (EuroSCORE II) and Society of Thoracic Surgeons (STS) risk-score for risk stratification in Indian patients undergoing cardiac surgery. *Ann Card Anaesth* 2013;16:163–6.
- [21] Özkan S, Özdemir F, Uğur O, Demirtunç R, Balci AY, Kizilay M, et al. The effects of the metabolic syndrome on coronary artery bypass grafting surgery. *Cardiovasc J Afr* 2017;28:48–53.
- [22] Ozyazicioglu A, Yalcinkaya S, Vural AH, Yumun G, Bozkurt O. Effects of metabolic syndrome on early mortality and morbidity in coronary artery bypass graft patients. *J Int Med Res* 2010;38:202–7.
- [23] Zapata D, Halkos M, Binongo J, Puskas J, Guyton R, Lattouf O. Effects and outcomes of cardiac surgery in patients with cardiometabolic syndrome. *J Card Surg* 2020;35:794–800.
- [24] Ardeshiri M, Faritus Z, Ojaghi-Haghighi Z, Bakhshandeh H, Kargar F, Aghili R. Impact of metabolic syndrome on mortality and morbidity after coronary artery bypass grafting surgery. *Res Cardiovasc Med* 2014;3:1–7.
- [25] Pimenta E, Passarelli Jr O, Borelli F, Sousa MG, Gun C, Amato V, et al. Metabolic syndrome in patients undergoing coronary artery bypass graft: prevalence and a marker of morbidity/mortality

- during hospitalization and 30 days after hospital discharge. *Arq Bras Cardiol* 2007;88:413–7.
- [26] Bugra AK, Göde S, Bugra A, Eltutan S, Arafat Z, Şen O, et al. Mediastinitis after cardiac surgery: risk factors and our vacuum-assisted closure results. *Kardiochir Torakochirurgia Pol* 2021;18:195–202.
- [27] Cappabianca G, Paparella D, Visicchio G, Capone G, Lionetti G, Numis F, et al. Preoperative C-reactive protein predicts mid-term outcome after cardiac surgery. *Ann Thorac Surg* 2006;82:2170–8.
- [28] Elenbaas TWO, Soliman Hamad MA, Schonberger JPAM, Martens EJ, van Zundert AAJ, van Straten AHM. Preoperative atrial fibrillation and elevated C-reactive protein levels as predictors of mediastinitis after coronary artery bypass grafting. *Ann Thorac Surg* 2010;89:704–9.
- [29] Tschudin-Sutter S, Meinke R, Schuhmacher H, Dangel M, Eckstein F, Reuthebuch O, et al. Drainage days – an independent risk factor for serious sternal wound infections after cardiac surgery: a case control study. *Am J Infect Control* 2013;41:1264–7.
- [30] Eton V, Sinyavskaya L, Langlois Y, Morin JF, Suissa S, Brassard P. Effect of pre-operative use of medications on the risk of surgical site infections in patients undergoing cardiac surgery. *Surg Infect* 2016;17:557–62.
- [31] Kayani WT, Banteali SJ, Lee VV, Elayda M, Khan A, Nambi V, et al. Association between statins and infections after coronary artery bypass grafting. *Int J Cardiol* 2013;168:117–20.
- [32] Oddsson SJ, Sigurdsson MI, Helgadóttir S, Sigurjonsson H, Viktorsson S, Arnorsson T, et al. Lower mortality following coronary arterial revascularization in patients taking statins. *Scand Cardiovasc J* 2012;46:353–8.
- [33] Young S, Young LK, Yong SC, Jong CK, Sang BH, Jae KS. Effect of preoperative statin therapy on myocardial protection and morbidity endpoints following off-pump coronary bypass surgery in patients with elevated C-reactive protein level. *Kor J Anesthesiol* 2010;58:136–41.
- [34] Boccara F, Cohen A, Di Angelantonio E, Meuleman C, Ederhy S, Dufaitre G, et al. Coronary artery bypass graft in HIV-infected patients: a multicenter case control study. *Curr HIV Res* 2008;6:59–64.
- [35] Dominici C, Chello M. Impact of human immunodeficiency virus (HIV) infection in patients undergoing cardiac surgery: a systematic review. *Rev Cardiovasc Med* 2020;21:411–8.
- [36] Jimenez-Exposito MJ, Mestres CA, Claramonte X, Cartana R, Josa M, Pomar JL, et al. Mortality and morbidity in HIV-infected patients undergoing coronary artery bypass surgery: a case control study. *Rev Espanol Cardiol* 2006;59:276–9.
- [37] Robich MP, Schiltz N, Johnston DR, Mick S, Tse W, Koch C, et al. Outcomes of patients with human immunodeficiency virus infection undergoing cardiovascular surgery in the United States. *J Thorac Cardiovasc Surg* 2014;148:3066–75.
- [38] Bortz J, Guariglia A, Klaric L, Tang D, Ward P, Geer M, et al. Biological age estimation using circulating blood biomarkers. *Commun Biol* 2023;6:1089.
- [39] National Institute for Health and Care Excellence. Routine preoperative tests for elective surgery. NICE guideline NG45. London: NICE; 2016. Available at: <https://www.nice.org.uk/guidance/ng45> [last accessed November 2024].
- [40] Boyle M, Vaja R, Rochon M, Luhana S, Gopalaswamy M, Bhudia S, et al. Sex differences in surgical site infections following coronary artery bypass grafting: a retrospective observational study. *J Hosp Infect* 2024;146:52–8.
- [41] Theodore DA, Goodwin RD, Zhang Y, Schneider N, Gordon RJ. History of depression and increased risk of sternal wound infection after cardiothoracic surgery: a novel and potentially modifiable risk factor. *Open Forum Infect Dis* 2019;6:ofz083.
- [42] Togan T, Gunday M, Ciftci O, Bingol H. Can preoperative erythrocyte sedimentation rate serve as an indicator for midterm adverse events after coronary bypass grafting? *Heart Surg Forum* 2015;18:E47–52.
- [43] Immohr MB, Sugimura Y, Aubin H, Rellecke P, Boeken U, Lichtenberg A, et al. Iron deficiency does not impair the outcome after elective coronary artery bypass and aortic valve procedures. *J Card Surg* 2021;36:542–50.
- [44] Suehiro K, Tanaka K, Matsuura T, Funao T, Yamada T, Mori T, et al. Preoperative hydroperoxide concentrations are associated with a risk of postoperative complications after cardiac surgery. *Anaesth Intensive Care* 2014;42:487–94.
- [45] Duarte JC, Reyes P, Bermudez D, Alzate JP, Maldonado JD, Cortes JA. Bacteriuria is not associated with surgical site infection in patients undergoing cardiovascular surgery. *Am J Infect Control* 2018;46:180–5.
- [46] Filsoufi F, Castillo JG, Rahmanian PB, Broumand SR, Silvay G, Carpentier A, et al. Epidemiology of deep sternal wound infection in cardiac surgery. *J Cardiothor Vasc Anesth* 2009;23:488–94.
- [47] Toumpoulis IK, Anagnostopoulos CE, DeRose Jr JJ, Swistel DG. The impact of deep sternal wound infection on long-term survival after coronary artery bypass grafting. *Chest* 2005;127:464–71.
- [48] Lemus Barrios GA, Cardenas Castellanos JM, Curcio Borrero CL, Moreno Gomez GA. Effects of frailty on the adverse outcomes of cardiac surgery in the elderly. *Rev Colomb Cardiol* 2020;27:250–61.
- [49] Bäck C, Hornum M, Olsen PS, Møller CH. 30-day mortality in frail patients undergoing cardiac surgery: the results of the Frailty in Cardiac Surgery (FICS) Copenhagen Study. *Scand Cardiovasc J* 2019;53:348–54.
- [50] Crape BL, Gusmanov A, Orazumbekova B, Davtyan K. Higher surgery and recovery room air pressures associated with reduced surgical site infection risk. *World J Surg* 2021;45:1088–95.
- [51] Hosseinrezaei H, Rafiei Amiri M. Incidence and risk factors of sternal wound infection at site of incision after open-heart surgery. *J Wound Care* 2012;21:408–11.
- [52] Jaimes MC, Torrado LAA, Reyes NFS, Mackenzie JC, Mallarino JPU. Hypothyroidism is a risk factor for atrial fibrillation after coronary artery bypass graft. *Braz J Cardiovasc Surg* 2017;32:475–80.
- [53] Kiriya Y, Toshiaki N, Shibasaki I, Ogata K, Ogawa H, Takei Y, et al. Sarcopenia assessed by the quantity and quality of skeletal muscle is a prognostic factor for patients undergoing cardiac surgery. *Surg Today* 2020;50:895–904.
- [54] Tadros MA, Williams VR, Plourde S, Callery S, Simor AE, Vearncombe M. Risk factors for *Staphylococcus aureus* surgical site infection during an outbreak in patients undergoing cardiovascular surgery. *Am J Infect Control* 2013;41:509–12.
- [55] Hassan M, Smith JM, Engel AM. Predictors and outcomes of sternal wound complications in patients after coronary artery bypass graft surgery. *Am Surg* 2006;72:515–20.
- [56] Maillet JM, Oghina G, le Besnerais P, Thierry S, Bouquet G, Mesnildrey P, et al. Preoperative carriage and postoperative same-species sternal wound infection after cardiac surgery. *Interact Cardiovasc Thorac Surg* 2011;13:381–5.
- [57] Munoz P, Hortal J, Giannella M, Barrio JM, Rodriguez-Creixems M, Perez MJ, et al. Nasal carriage of *S. aureus* increases the risk of surgical site infection after major heart surgery. *J Hosp Infect* 2008;68:25–31.
- [58] Cutrell JB, Barros N, McBroom M, Luby J, Minhajuddin A, Ring WS, et al. Risk factors for deep sternal wound infection after cardiac surgery: influence of red blood cell transfusions and chronic infection. *Am J Infect Control* 2016;44:1302–9.
- [59] Dodds Ashley ES, Carroll DN, Engemann JJ, Harris AD, Fowler Jr VG, Sexton DJ, et al. Risk factors for postoperative mediastinitis due to methicillin-resistant *Staphylococcus aureus*. *Clin Infect Dis* 2004;38:1555–60.
- [60] Mehaffey JH, Hawkins RB, Charles EJ, Thibault D, Williams ML, Brennan M, et al. Distressed communities are associated with

- worse outcomes after coronary artery bypass surgery. *J Thorac Cardiovasc Surg* 2020;160:425–32.
- [61] Al Salmi H, Elmahrouk A, Arafat AA, Edrees A, Alshehri M, Wali G, et al. Implementation of an evidence-based practice to decrease surgical site infection after coronary artery bypass grafting. *J Int Med Res* 2019;47:3491–501.
- [62] Abdelnoor M, Vengen OA, Johansen O, Sandven I, Abdelnoor AM. Latitude of the study place and age of the patient are associated with incidence of mediastinitis and microbiology in open-heart surgery: a systematic review and meta-analysis. *Clin Epidemiol* 2016;8:151–63.
- [63] Benedetto U, Dimagli A, Gibbison B, Sinha S, Pufulete M, Fudulu D, et al. Disparity in clinical outcomes after cardiac surgery between private and public (NHS) payers in England. *Lancet Reg Health Eur* 2021;1:100003.
- [64] van Laar C, Timman ST, Noyez L. Decreased physical activity is a predictor for a complicated recovery post cardiac surgery. *Health Qual Life Outcomes* 2017;15:5.
- [65] Robinson PJ, Billah B, Leder K, Reid CM. Factors associated with deep sternal wound infection and haemorrhage following cardiac surgery in Victoria. *Interact Cardiovasc Thorac Surg* 2007;6:167–71.
- [66] Cayci C, Russo M, Cheema F, Martens T, Ozcan V, Argenziano M, et al. Risk analysis of deep sternal wound infections and their impact on long-term survival: a propensity analysis. *Ann Plast Surg* 2008;61:294–301.
- [67] Nešpor D, Fabián J, Némec P. A retrospective analysis of deep sternal wound infections after longitudinal median sternotomy. *Cor Vasa* 2015;57:e75–81.
- [68] Tully PJ, Cardinal T, Bennetts JS, Baker RA. Selective serotonin reuptake inhibitors, venlafaxine and duloxetine are associated with in hospital morbidity but not bleeding or late mortality after coronary artery bypass graft surgery. *Heart Lung Circ* 2012;21:206–14.
- [69] Batista R, Kaye K, Yokoe DS. Admission-specific chronic disease scores as alternative predictors of surgical site infection for patients undergoing coronary artery bypass graft surgery. *Infect Control Hosp Epidemiol* 2006;27:802–8.
- [70] Kalyoncuoglu MaO S, Sahin M. Does CHA<sub>2</sub>DS<sub>2</sub>-VASC score predict MACE in patients undergoing isolated coronary artery bypass grafting surgery? *Braz J Cardiovasc Surg* 2019;34:542–9.
- [71] Hayashi J, Uchida T, Ri S, Hamasaki A, Kuroda Y, Yamashita A, et al. Clinical significance of the prognostic nutritional index in patients undergoing cardiovascular surgery. *Gen Thorac Cardiovasc Surg* 2020;68:774–9.
- [72] Lee SI, Ko KP, Choi CH, Park CH, Park KY, Son KH. Does the prognostic nutritional index have a predictive role in the outcomes of adult cardiac surgery? *J Thorac Cardiovasc Surg* 2020;160:145–53.