

# Deep sternal wound infection after cardiac surgery in the Chinese population: a single-centre 15-year retrospective study

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**Background:** Deep sternal wound infection (DSWI) is a severe complication following cardiac surgery. A retrospective study was implemented to determine the risks and clinical characteristics of DSWI after cardiac operation in the Chinese population.

**Methods:** We analysed 7,944 consecutive patients who underwent cardiac surgery via median sternotomy from January 2002 to December 2016 at our institution. Multiple logistic regression analysis was used to identify risk factors for DSWI.

**Results:** A total of 106 patients (1.33%) suffered from DSWI; significant risk factors included body mass index (BMI) ( $P=0.02$ ; OR=1.08; 95% CI: 1.01–1.16) and reoperation ( $P<0.01$ ; OR=5.93; 95% CI: 2.88–12.25). The most common bacterium involved in DSWI was *staphylococcus aureus* (23%). Among all DSWI patients, the overall survival rate of the group treated with flap reconstruction was significantly higher than that in the group treated with intravenous antibiotics and sternal debridement (87% vs. 59%,  $P=0.01$ ).

**Conclusions:** DSWI was associated with several risk factors. Effective intervention strategies could improve the outcome of patients undergoing cardiac surgery.

**Keywords:** Deep sternal wound infection (DSWI); median sternotomy; wound infection; mediastinitis

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## Introduction

Median sternotomy has been the most frequently used incision for cardiac surgery since its introduction in 1956 (1). One of the life-threatening complications that occurs in patients following cardiovascular surgery through median sternotomy is deep sternal wound infection (DSWI). The incidence of DSWI that has been reported ranges from 0.2% to 8% (2,3). However, the mortality varies from 8% to 45%. Additionally, DSWI is associated with increased medical cost, prolonged hospital stays and even reoperation (4–6). Therefore, precautionary intervention before cardiac

surgery and effective treatment after the occurrence of DSWI are significant.

Previous studies have attempted to determine risk factors of DSWI. However, a consensus on the significant risk factors of DSWI has not been reached. The identified risk factors include obesity, diabetes mellitus, tracheostomy, use of the bilateral mammary arteries, reoperation, transfusion, and prolonged operation time (6,7). Patients who suffer from DSWI is divided into three types according to time between operation and onset of DSWI, each of type has different pathogenic bacteria. *Staphylococcus* was the most common pathogenic bacterium in DSWI patients (8).

**Table 1** Definition of DSWI

DSWI requires the presence of one of the following criteria
Positive culture results of mediastinal tissue or fluid
Mediastinitis was observed during the operation
The presence of either chest pain, sternal instability or fever (>38 °C) and purulent drainage from the drained placed beneath the fascial layer, or an organism present in a blood culture
DSWI, deep sternal wound infection.

**Table 2** Type of DSWI

Type	Description
I	DSWI presenting within 1 weeks after the operation
II	DSWI presenting within 2–4 weeks after the operation
III	DSWI presenting within months to years after the operation
DSWI, deep sternal wound infection.	

Although studies of DSWI have been reported for many years, little had been shown about DSWI in the Chinese population. The present study was designed to investigate the clinical characteristic of DSWI in patients who underwent cardiac surgery at our institution in order to conduct timely effective intervention.

## Methods

### Study design

A retrospective study was performed at the department of cardiothoracic surgery at the Drum Tower Hospital that is affiliated with the medical school of Nanjing University. Consecutive patients who underwent cardiac surgery via median sternotomy during 1 January 2002 and 31 December 2016 were included. All patients were Chinese. The patients were diagnosed with DSWI by an attending physician and an infection control physician during their hospital stay and readmission period based on the definition of DSWI (*Table 1*) (9,10). The time interval between the operation and the date of diagnosis was obtained to determine the type of DSWI (11) (*Table 2*). We have treated these DSWI patients 15 years ago and use old diagnoses criteria at that time. In order to ensure the validity and consistency of data, we select the old definition. So, patients who suffer from sternal wound infection between one week after operation are excluded.

### Patient selection

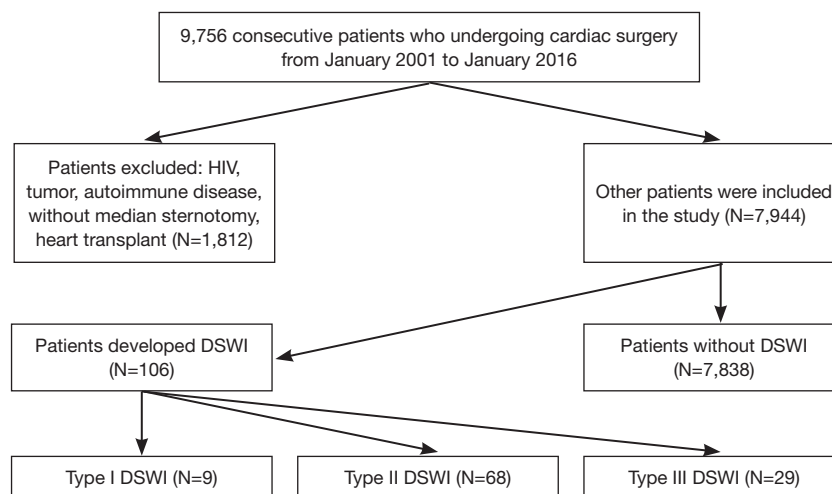
As follow chart showed, the study group did not include patients who underwent heart transplantation operations (*Figure 1*). Patients who suffered preoperative infection were divided into two types: patients who suffered from a simple bacterial infection were included into the study after antibiotic treatment (pneumonia, infective endocarditis and urinary tract infection); and patients who suffered from immune diseases were excluded from the study (HIV, tumour and autoimmune diseases). To confirm the diagnosis of DSWI, the group reviewed the database and collected relevant information regarding the patients, including their clinical records and laboratory examination results. Control patients were selected by the study group among eligible patients without evidence of DSWI or superficial wound infection. Patients who had delayed sternal closure after cardiac surgery were excluded from both the case and control patient groups.

### Operation procedure

Antibiotic prophylaxis was used in all patients; cefuroxime was most commonly used, starting 30 minutes before the operation until 72 hours after surgery. The interlocked figure of “8” wire closure technique was generally used in patients who underwent cardiac surgery through a median sternotomy. However, if patients had an osteoporotic sternum, the Robicsek technique was adopted. Establishment of cardiopulmonary bypass (CPB) involved the vena cava and aorta or vena cava, axillary artery and femoral artery. All surgeries were performed by four experienced surgeons at the institution.

### Treatment of DSWI

Treatment of DSWI included antibiotics, sternal debridement and flap reconstruction. Cefuroxime was the most commonly used agent. However, if patients were allergic to cefuroxime or culture MASA after the operation, cefuroxime was replaced by vancomycin. Tienam was added to the antibiotic treatment in patients with complicated respiratory diseases. The pectoral muscle flap was the most commonly used material for flap reconstruction. Extensive wound debridement was the first step of flap reconstruction, which involved resecting the necrotic tissue and sternum. Then, the tissue defect caused by the resection was closed by the transposition of the pectoralis major flap. In patients who only suffered from a minor tissue defect, iodine irrigation was performed instead.



**Figure 1** Patient selection. DSWI, deep sternal wound infection.

**Table 3** The demographics and operative variables of eligible patients.

Variables	DSWI group (N=106)	Control group (N=7,838)	P
Age (y)	56±1	51±35	0.01
Male	58 (55%)	4,222 (54%)	0.86
Diabetes	12 (11%)	525 (7%)	0.06
Hypertension	29 (27%)	1,657 (21%)	0.12
BMI (kg/m <sup>2</sup> )	25±3	23±4	<0.01
Isolated CABG procedure	28 (26%)	1,157 (15%)	<0.01
Isolated valve procedure	33 (31%)	3,324 (42%)	<0.01
CABG + valve procedure	11 (10%)	499 (6%)	0.09
Aortic procedure	16 (15%)	1,251 (16%)	0.81
CPB time (min)	174±16	158±9	0.07
Transfusion	47 (44%)	3,207 (41%)	0.48
Reoperation	11 (10%)	202 (3%)	<0.01

Isolated CABG procedure indicates patients who only underwent the CABG procedure. Isolated valve procedure indicates patients who only underwent a valve procedure. DSWI, deep sternal wound infection; BMI, body mass index; CABG, cardiac artery bypass graft procedure; CPB, cardiopulmonary bypass.

### Statistical analysis

Comparisons of the categorical variables were performed using the chi-square test. Comparisons of the continuous variables were performed by one-way ANOVA. The multiple

logistic regression was used to identify the most significant mortality predictors. After the univariate analysis, risk factors with a P value <0.1 were selected for the multivariate model. Statistical significance was assessed at  $\alpha=0.05$ . All analyses were performed by the SPSS version 17 software.

### Results

Table 3 shows the patient characteristics and operation data. There were 7,944 patients included in the study between January 2002 and December 2016, among whom 106 patients met the criteria of DSWI, and 7,838 patients were included as control patients. Several variables were different between the case patients and the control patients, including age ( $P=0.01$ ), diabetes ( $P=0.06$ ), body mass index (BMI) ( $P<0.01$ ), isolated cardiac artery bypass graft procedure (CABG) ( $P<0.01$ ), isolated valve procedure ( $P<0.01$ ), CPB time ( $P=0.07$ ) and reoperation ( $P<0.01$ ). Then, we conducted a multiple logistic regression analysis that included age, BMI, CABG, CABG with valve surgery, CPB time, transfusion and reoperation as risk factors. The results are summarized in Table 4. BMI (OR=1.08;  $P=0.02$ ; 95% CI: 1.01–1.16) and reoperation (OR=5.93;  $P<0.01$ ; 95% CI: 2.88–12.25) were identified as significant risk factors for DSWI. In particular, patients who suffered from reoperation were five times as likely to have DSWI compared with patients who did not. Although BMI is not a risk factor of DSWI, we observed that the association of BMI and outcomes is inverted “V” shape (Figure S1).

All patients who were suspected of having DSWI or

had symptoms of DSWI underwent a blood culture and an exudation culture. It is worth noting that the overall number of positive bacterial culture was observed in only 38 out of 106 patients (35.85%). We identified that the

**Table 4** Multivariate conditional logistic regression results for DSWI

Variables	P	Odds ratio	95% confidence interval
Age	0.81	0.99	0.98–1.02
BMI	0.02	1.08	1.01–1.16
Isolated CABG	0.38	2.51	0.32–19.41
CABG + valve procedure	0.08	1.89	0.92–3.87
CPB time	0.06	1.01	1.00–1.01
Transfusion	0.24	1.35	0.82–2.20
Reoperation	<0.01	5.93	2.88–12.25

DSWI, deep sternal wound infection; BMI, body mass index; CABG, cardiac artery bypass graft procedure; CPB, cardiopulmonary bypass.

**Table 5** Pathogenic data of the DSWI patients

Aetiology	Patients with a diagnosis of aetiology [n (%)]
GNB	13 (34%)
Pseudomonas aeruginosa	3 (8%)
Acinetobacter baumannii	8 (21%)
Enterobacter cloacae	2 (5%)
GPC	14 (37%)
Staphylococcus aureus	9 (24%)
Methicillin resistant staphylococcus aureus (MRSA)	2 (5%)
Enterococcus faecalis	3 (8%)
Mixed infection	11 (29%)

DSWI, deep sternal wound infection; GNB, gram-negative bacteria; GPC, gram-positive bacteria.

**Table 6** The influence of different treatments on the survival rate

Infection type	Debridement + flap reconstruction (N=30)	Debridement (N=76)	P	OR (95% CI)
I	0/1 (0%)	6/8 (75%)	0.44	0.29 (0.04–2.45)
II	14/15 (93%)	34/53 (64%)	0.07	0.43 (0.18–1.03)
III	12/14 (86%)	5/15 (33%)	0.01	3.56 (1.43–8.87)

OR, odds ratio; CI, confidence interval.

most common bacteria that were isolated from the wound were *staphylococcus aureus* (n=9, 23.68%) and *Acinetobacter baumannii* (n=8, 21.05%) in patients diagnosed DSWI, as shown in *Table 5*. The positive rate of gram-positive organisms (n=14, 36.84%) was comparable with the positive rate of gram-negative organisms (n=13, 34.21%). In addition, there was mixed infection in patients with DSWI (n=11, 28.95%).

Based in the time interval between the operation and the date of DSWI diagnosis, patients in the DSWI group were divided into three types of DSWI, as summarized in *Table 6*. Patients who were diagnosed as type III exhibited a significant difference between the two treatment groups (P=0.01). But, we do not observed difference in diabetes patients (*Tables S1-S3*). Compared with antibiotic treatment and debridement, which were widely used in patients with DSWI, flap reconstruction was reported as an effective treatment in those patients. Hence, we analysed the survival rate of patients who were treated with antibiotics, debridement and flap reconstruction. As showed in *Table 6*, the latter group had an obviously better prognosis than the former group. It may be worth indicating that timely surgical intervention was better than conservative treatment in patients with DSWI, especially in patients who were diagnosed as type III DSWI (P=0.01).

## Discussion

In the present study, we conducted a retrospective study on patients who underwent a cardiac operation in the past 15 years. To our knowledge, this is the largest retrospective study about DSWI in the Chinese population. Recently, some definition of criteria of DSWI have changed. In order to ensure the validity and consistency of data, we select the old definition. But new definition of DSWI is more suitable to current clinic status (10). The investigation provided an overall evaluation of DSWI after cardiac surgery in the Chinese population. Robinson et al reported that the incidence of DSWI was 1.3% in 12,000 patients who

underwent cardiac surgery from July 2001 to June 2005 (6). We found that the incidence of DSWI is 1.33% and is comparable with the report above. However, considering that there were case patients who were not readmitted to the institution, the occurrence of DSWI may have been underestimated. Improvements in aseptic technique and the prevention of antibiotic use may contribute to a reduction in the incidence of DSWI after cardiac surgery. Some studies have reported that antibiotic prophylaxis was beneficial for reducing the incidence of DSWI after the operation (12-15).

The occurrence of DSWI has multifactorial causes. Floros *et al.* determined that the risk factors for DSWI consisted of smoking, re-exploration, oral hypoglycemic drug, emergency surgery, obesity, IABP and transfusion (2). In addition, Pilarczyk *et al.* identified that percutaneous dilatational tracheotomy within 48 hours after cardiac surgery was associated with mediastinitis (16). Our study identified that the potential risk factors for DSWI were BMI and reoperation. Although it was not identified as a significant risk factor, age was a significant variable between case patients and control patients, as the study group was older than the control group. Patients who have higher BMI are in danger of sternal instability. Therefore, sternal instability may contribute to the development of DSWI in elderly patients who received cardiac surgery via a median sternotomy.

Furthermore, our results presented that reoperation was a risk factor for DSWI after operation and that CPB time has a trend of difference ( $P=0.06$ ). It was a little different from other studies that have been published (17-21), which have reported that operation time but not CPB time was a risk factor for DSWI. Prolonged CPB time may be associated with complex procedures and more surgical implants; patients who undergo reoperation often suffer a worse tissue situation and need more time, usually because of bleeding or unstable haemodynamics. These situations probably significantly increase DSWI risk, but an additional randomized control trial is needed to verify the conjecture (2,20,22,23). Although transfusion rates were not significantly different in our study, there was a trend of difference. Whether there are other factors involved in DSWI in patients included in our study population need further investigation.

Pathogenic microorganisms were confirmed with exudation culture. The low rate of positive results may be attributed to: (I) the selection pressure of antibiotic prophylaxis, as broad spectrum antibiotic is routinely used before cardiac operation at present, which may result in a

negative result from blood culture and exudation culture; (II) aetiology of DSWI was atypical pathogens (mycoplasma, anaerobic bacteria), which are difficult to culture *in vitro*; (III) there were some DSWI patients who underwent sternal dehiscence without infection; and (IV) antibiotic abuse is still a complicated problem in the treatment of infection in the Chinese population. The most common bacterium was staphylococcus aureus, which was in agreement with a previous study. However, the proportion of staphylococcus aureus varied from previous reports (2,8,21,24,25). Some studies have attempted to investigate the benefit of antibiotic prophylaxis and confirm that intervention strategies were efficient (12,13,22,26).

For the treatment of DSWI, it is thought that debridement was the most effective treatment after a timely diagnosis, and the use of antibiotics targeting the pathogenic bacterium was better than dilute povidone-iodine irrigation for the treatment of DSWI (23,27). Some studies have reported that an antibiotic-collagen sponge was effective for the prevention of DSWI. However, the results from different studies have been controversial (12,13,28,29). In this study, we showed that the group treated with intravenous antibiotic, sternal debridement and flap reconstruction had a better prognosis than the group treated with intravenous antibiotic and sternal debridement (30-33). Negative pressure wound therapy system is also used in the treatment of DSWI patients. But, due to this treatment equipment is applied in our department recently, not of all patients received this new therapy system. So, the influence of negative pressure wound therapy system on DSWI patients remain to be further study. The periods are not consecutive. Someone with an infection at 10 days is not included in either type I or type II. It is noteworthy that the proportion of type III DSWI patients are significantly different between the two groups. Considering this, we assumed that the type of DSWI may a considerable factor that affects the results of treatment. In died patients, time to death after diagnosis of infection is respectively  $55\pm 14$  and  $34\pm 3$  days in debridement and flap reconstruction group and debridement group. So, if they died before they were well enough to get a flap reconstruction included in the debridement group, there, there exist potential bias. Overall, the results indicate that timely surgical intervention is beneficial for patients with DSWI (23).

Although the findings were statistically supported with a large sample size, there were some limitations that may have affect the findings. First, this is a retrospective study, which cannot account for the effect of unknown variables.

In addition, the low event incidence, such as risk factors and bacteriological characteristics, precluded us from further investigation about the details of DSWI. Furthermore, the positive rate of pathogenic bacteria was low and may have affected the result. Therefore, a prospective randomized controlled trial should be implemented to analyse the characteristics of DSWI in the Chinese population.

## Conclusions

In this retrospective study of DSWI in the Chinese population, we determined the incidence of DSWI in patients who underwent cardiac surgery via median sternotomy and the pathogenic characteristics of the bacteria. DSWI was associated with several risk factors, including BMI and reoperation. Additionally, flap reconstruction is a valid treatment for patients with DSWI, especially for patients with type III DSWI. Effective intervention strategies could improve the outcome of patients undergoing cardiac surgery

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## Footnote

*Conflicts of Interest:* The authors have no conflicts of interest to declare.

*Ethical Statement:* The study was approved by the institutional ethics board of Nanjing Drum Tower Hospital (No. 2017-090-01).

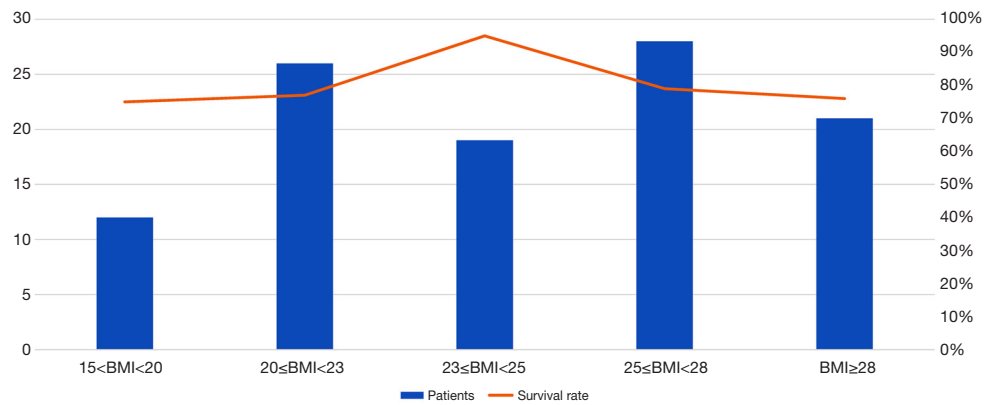
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## Supplementary



**Figure S1** The association between BMI and survival rate of DSWI. BMI, body mass index; DSWI, deep sternal wound infection.

**Table S1** The demographics and operative variables of diabetes patients

Variables	DSWI group (N=12)	Control group (N=525)	P
Age (y)	69±7	52±17	<0.01
Male	7 (58%)	282 (54%)	0.75
Hypertension	10 (83%)	348 (66%)	0.35
BMI (kg/m <sup>2</sup> )	27±3	23±4	<0.01
Isolated CABG procedure	8 (67%)	71 (14%)	<0.01
CPB time (min)	187±38	159±67	0.49
Transfusion	6 (50%)	221 (42%)	0.58

Isolated CABG procedure indicates patients who only underwent the CABG procedure. Isolated valve procedure indicates patients who only underwent a valve procedure. DSWI, deep sternal wound infection; BMI, body mass index; CABG, cardiac artery bypass graft procedure; CPB, cardiopulmonary bypass.

**Table S2** The influence of different treatments on the survival rate in diabetes patients

	Debridement + flap reconstruction (N=6)	Debridement (N=6)
Infection type		
I	1 (17%)	0
II	3 (50%)	6 (100%)
III	2 (33%)	0
Survival rate	5 (83%)	4 (67%)

**Table S3** The influence of different treatments on the survival rate

Different treatments	Diabetes	Non-diabetes	P
Debridement	6	70	0.15
Debridement + flap reconstruction	6	24	