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Sternal Wound Complications Following Cardiac Surgery

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1. Introduction

Median sternotomy is a commonly performed incision with distinct advantages for exposure of mediastinal and pulmonary hilar structures \cite{1}. However, a well-defined incidence of wound complications is associated with sternotomy, which are costly and potentially lethal in cases of deep sternal wound infection (DSWI) or mediastinitis \cite{2-13}. Not only is DSWI associated with significant perioperative mortality, but historically even successfully treated DSWI is associated with reduced mid- and long-term survival compared with matched cardiac surgical patients without this dreaded postoperative complication \cite{7-12}.

In the past 10 years, we have accumulated extensive experience with managing DSWI as a referral center for these difficult problems. We and others have formalized a protocol for managing mediastinal infection utilizing negative pressure wound therapy which allows sternal salvage and improved outcomes in the majority of cases of DSWI \cite{14-16}. This report describes our protocol for managing mediastinitis and presents our results for the past 18 years.

2. Background

Median sternotomy was originally introduced by Milton in 1897 and was performed infrequently for various conditions of the mediastinum until cardiac surgery as a field blossomed in the 1950s \cite{17}. Shumacker first suggested median sternotomy as the procedure of choice for approaches to the heart and great vessels \cite{18} since it avoided the significant pain and other complications, primarily pulmonary- or pleural-based, of the bilateral anterior thoracotomy (“clamshell”) incision, which was most frequently used up to that point. However, not until Julian and colleagues demonstrated discrete advantages of the median sternotomy incision for cardiac surgery, particularly improved surgical efficiency, excellent exposure the heart, great vessels, and pulmonary hila, and reduced pulmonary trauma, was a convincing argument for median sternotomy as the incision of choice for cardiac surgical procedures put forth \cite{19 Table 1}. The utility of this incision, the ease and speed of performance, and the common nature of surgically-treated cardiovascular and
<table>
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<tr>
<th>STRUCTURE</th>
<th>STERNOTOMY</th>
<th>RIGHT THORACOTOMY</th>
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<td>MAIN STEM</td>
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<td>BRONCHI</td>
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Table 1. Comparison of exposure of various intrathoracic anatomic structures through median sternotomy, right thoracotomy, or left thoracotomy. (+++ denotes excellent, reliable exposure; 0 denotes no reliable exposure)

thoracic diseases in multiple populations approached through this incision are factors which combine to make median sternotomy the most commonly performed osteotomy worldwide [20]. Recently, several alternatives to median sternotomy have been promoted, including thoracoscopic and robotic approaches to cardiac and thoracic procedures [21, 22]. These approaches are possible as a result of high fidelity instrumentation and video platforms but are limited by steep learning curves and expense. Therefore, for most hospital systems performing cardiac surgical procedures, median sternotomy remains the mainstay incision.

### 3. Definition and classification

There is still no universally accepted method for treatment of DSWI or other sternal wound complication. One possible explanation for this is that until relatively recently, thorough classification schemes providing specific criteria for assignment as deep sternal infection, mediastinitis, superficial infection, or sterile sternal dehiscence were lacking. However, it is important to distinguish between these individual entities since each demands a different management strategy. Furthermore, results of reported series which include a variety of wound complications will differ from those that report a homogenous population. For example, sternal wound infections that are limited to the superficial soft tissues obviously demand less aggressive intervention for treatment and will generally respond more readily to treatment than deep sternal infection with higher likelihood for success. Therefore, when
comparing different treatment modalities for sternal wound infection, it is important to be clear about the extent of infection, since heterogeneity could skew reported results.

Two prominent classification schemes have been proposed and are in use. Mediastinal dehiscence or the more chronic form, sternal nonunion, is defined as sternal wound disruption without any evidence for infection either clinically or pathologically [23, 24]. These entities will not be discussed in any detail. In contrast, mediastinitis, as characterized by the U.S. Centers for Disease Control and Prevention (CDC), is an infection of the mediastinum diagnosed by isolation of pathogenic organisms from the mediastinal fluid or tissue especially when there is obvious evidence of infection at the time of sternal exploration [25]. Alternatively, a combination of clinical features including chest pain, sternal drainage with bony instability, fevers, radiographic findings such as widened mediastinum, and bacterial isolation may also warrant a diagnosis of mediastinitis. Obviously, from the surgical perspective, these definitions are somewhat lacking since either superficial infections, confined to the soft tissues, or deep infections, involving the bone and/or retrosternal space could produce bacteremia and clinical signs of severe infection [23].

Fortunately, more descriptive classification schemes have been introduced and provide more specific insight into the pathologic involvement of the sternal tissues and the clinical consequences and course [23, 26, 27]. For example, the classification scheme introduced by El Oakley and Wright is based on the time at which the patient presents with mediastinitis relative to the initial surgical procedure [23]. Schulman et al have advocated a similar classification system [27]. In addition, the El Oakley description also accounts for relevant risk factors underlying the clinical scenario and whether or not previous attempts to treat the sternal wound infection have been made and failed [Table 2]. Therefore, five distinct categories of infection are described, each with important treatment implications. For example, the subtypes I and II appear to respond well to primary sternal closure with mediastinal irrigation, while subtypes III-V appear to require more aggressive sternal debridement and repair techniques [23].

<table>
<thead>
<tr>
<th>Classification</th>
<th>Description of Infection</th>
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<tbody>
<tr>
<td>Type 1</td>
<td>Mediastinitis within 2 weeks of operation without risk factors</td>
</tr>
<tr>
<td>Type 2</td>
<td>Mediastinitis between 2 and 6 weeks of operation without risk factors</td>
</tr>
<tr>
<td>Type 3a</td>
<td>Type 1 with one or more risk factors</td>
</tr>
<tr>
<td>Type 3b</td>
<td>Type 2 with one or more risk factors</td>
</tr>
<tr>
<td>Type 4a</td>
<td>Type 1, 2, or 3 after one failed therapy</td>
</tr>
<tr>
<td>Type 4b</td>
<td>Type 1, 2, or 3 after more than one failed therapy</td>
</tr>
<tr>
<td>Type 5</td>
<td>Mediastinitis presenting 6 weeks or more after operation</td>
</tr>
</tbody>
</table>

Table 2. Classification scheme of mediastinitis introduced by El Oakley and Wright based on the time at which the patient presents with mediastinitis relative to the initial surgical procedure (modified from El Oakley and Wright [23]).
The classification by Jones et al differs from that of El Oakley and Wright in that it is more descriptive anatomically and physiologically [26, TABLE 3]. Three different “types” of sternal infection are described, encompassing both superficial and deep infections, and based on the degree of underlying tissue involvement with infection. We have preferred the use of this classification system as it is simpler to use since it is based strictly on features observed or encountered at the time of initial sternal exploration. In addition, Type 3b is physiologically meaningful since it denotes the patient who is systemically ill from the sternal wound process. In our own institutional experience of 222 adult cardiac surgical patients treated for postoperative DSWI, approximately 50% of patients exhibited septicemia (Jones 3b) upon initial presentation [14].

<table>
<thead>
<tr>
<th>Class</th>
<th>Depth of Involvement</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>1a</td>
<td>Superficial</td>
<td>Skin/subcutaneous tissue dehiscence</td>
</tr>
<tr>
<td>1b</td>
<td>Superficial</td>
<td>Exposed deep fascia, sutures intact</td>
</tr>
<tr>
<td>2a</td>
<td>Deep</td>
<td>Exposed bone, stable wired sternotomy</td>
</tr>
<tr>
<td>2b</td>
<td>Deep</td>
<td>Exposed bone, unstable wired sternotomy</td>
</tr>
<tr>
<td>3a</td>
<td>Deep</td>
<td>Exposed necrotic or fractured bone, unstable, heart exposed</td>
</tr>
<tr>
<td>3b</td>
<td>Deep</td>
<td>Types 2 or 3 with septicemia</td>
</tr>
</tbody>
</table>

Table 3. Mediastinal wound classification system modified from Jones et al [26]. Although anatomic involvement by infection is distinguished, the presence of septicemia is the most important feature clinically.

4. Risk factors and pathophysiology

Multiple different etiologies for development of DSWI have been invoked over time. For example, most infections were traditionally thought to arise as a result of breaks in proper surgical technique, prompting strict guidelines for sterile surgical technique. In addition, secondary involvement the mediastinum from remote sites such as leg incisions or the pulmonary tree has also been suggested as a mechanism for DSWI. The “endogenous pathway,” seeding of the mediastinum from other host sources, does appear to be important in the development of *S. aureus* mediastinitis. To illustrate, Jakob et al showed that nasal carriage of *S. aureus* was an independent predictor of sternal infection postoperatively [28], while others have demonstrated that application of mupirocin to the nares of *S. aureus* colonized individuals can help reduce postoperative infection [29]. More recently, other factors have been recognized as important contributors to cardiac surgical site infections. For instance, the appropriate timing of perioperative antibiotics and strict perioperative glucose control are both associated with reduced surgical wound infectious complications [30, 31].

Several different patient-related factors have been repeatedly implicated in the development of DSWI, most consistently obesity and diabetes mellitus [5, 6, 32, TABLE 4]. Eklund et al and others have observed that increasing severity of obesity elevates the risk for surgical site infection in a step-wise fashion [6, 13, 33]. Furthermore, when obesity is described in terms of percentage body fat, the relationship between obesity and surgical site infection is nearly linear and appears more accurate than use of body mass index as a descriptor of obesity [34]. Potential explanations for relationship between obesity and DSWI include technical problems.
Table 4. Compiled analyses for underlying risk factors associated with sternal wound infection. Obesity and diabetes mellitus are consistently shown to be independent predictors of poststernotomy mediastinitis.

<table>
<thead>
<tr>
<th>Study</th>
<th># Patients</th>
<th>Identified Risk Factors</th>
</tr>
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<tbody>
<tr>
<td>Cayci C, et al. Ann Plast Surg 2008 [52]</td>
<td>7,978</td>
<td>Obesity (BMI&gt;30 kg/m²), diabetes, urgent operation, recent smoking; h/o CVA, total HLOS, sepsis/endocarditis postop</td>
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during surgery owing to the patient’s size, increased bleeding, increased deadspace in the wound, and ineffective or inadequate dosing of perioperative antibiotics [6, 9]. Although multiple mechanisms for sternal wound complications are proposed, it is widely accepted that reduced sternal perfusion, often by virtue of internal mammary artery (IMA) harvesting for use as a vascular conduit in coronary artery revascularization, is one of the most important causes of sternal nonhealing and infection [1-6, 8, 26], especially when both IMAs are harvested for bypass graft surgery [5]. Therefore, more cases of DSWI appear to occur after coronary artery bypass grafting or after combined procedures that include coronary artery surgery [7, 35]. Other viable explanations for DSWI etiology include poor bone stock from osteoporosis, malnutrition, and other factors; poorly performed sternotomy leading to sternal fractures and/or costosternal disassociation; and other patient related factors including peripheral vascular disease and lung disease [8, 36]. Several other factors have been implicated in the development of DSWI but may not be manifest in the context of a retrospective review or randomized trial because the numbers are too low. For example, it is generally accepted that postoperative steroids or chronic immunosuppression increases risk for DSWI, but this has been difficult to demonstrate in even large database reviews [13, 37]. Finally, Risnes et al have demonstrated in a review of over 18,000 consecutive patients undergoing coronary artery surgery in Norway, the major preventable risk factor associated with the development of DSWI was the amount of blood product transfusion perioperatively [12].
Several risk analyses are available to estimate the individual patient’s propensity for developing DSWI. For example, using Society of Thoracic Surgery National Cardiac Database information, Fowler et al. created a model to estimate the risk for systemic infection after coronary artery bypass surgery using patient characteristics available preoperatively [13]. The Fowler model also provides for inclusion of important intraoperative details known to influence postoperative infection including the need for intra-aortic balloon counterpulsation and prolonged cardiopulmonary bypass times [13]. Although the model devised by Fowler et al. was based on various cases of major infection after coronary artery surgery, including DSWI, the authors did validate the model as predictive of infection in a test population from the STS Database, and the model was also recently validated in a different cohort of patients from the UK as being predictive of DSWI [38]. The EuroSCORE system also has been shown to predict infection and associated mortality with acceptable discrimination [39].

5. Incidence

Despite advancements in most aspects of perioperative care, rates of sternal wound complications, including mediastinitis, following adult cardiac surgery have varied little over the past 30 years [2-6], although incremental reduction in single center rates of DSWI have been noted and attributed to adjunctive measures such as strict perioperative glucose control [40]. As noted previously, rates of DSWI vary with regard to the specific definition used to describe the pathology encountered and the patient population studied. In general, reported rates of DSWI are relatively low, ranging from 0.25% to 3.6% [2-9, 38, 39, 41, 42]. However, the incidence of superficial sternal infection exceeds that of deep sternal infection by as many as three times [10, 41, 43], and Francel has stated that as many as 70% of patients with poststernotomy infection will have superficial tissue involvement [44].

6. Diagnosis

Depending on the virulent nature of the sternal wound infection, patients typically present within 30 days of cardiac surgery. The most common symptoms include sternal wound drainage, sternal instability, fever, and malaise. In many cases, a high index of suspicion is required to establish the diagnosis, especially when classic signs and symptoms are absent. In addition, it is often difficult to distinguish on physical examination the difference between deep sternal wound infection (infection present beneath the sternum) and superficial sternal infection. Therefore, early wound opening and inspection with appropriate sampling of tissue for bacteriologic assessment is strongly encouraged when sufficient clinical suspicion exists.

One of the most reliable signs of DSWI is sternal instability, which frequently implies a deeper problem than can be appreciated at the superficial level [44]. While sternal instability can be tested on physical examination, we have often noted that the patient themselves will typically report sternal clicking with inspiration, cough, or various other physical maneuvers, so this information should be thoughtfully considered if DSWI diagnosis is being entertained.

The use of various radiographic examinations is often encouraged when evaluating for DSWI, and, in fact, is included as part of the CDC guidelines for defining mediastinitis [25]. Formerly, PA and lateral chest radiograph was the investigative procedure of choice, where details such
as a “sternal stripe” indicated present of air between the two sternal halves. The lateral displacement of one or more sternal wires, secondary to tearing of the wire through one side of the sternum, has been a frequently noted finding in the case of poststernotomy infection [37]. More recently, chest computed tomography has been suggested as the procedure of choice for assessing sternal wound infection when a diagnosis cannot be established by clinical examination alone [45, 46]. Mediastinal fluid collections, free gas bubbles, soft tissue swelling, pleural effusions, sternal dehiscence, and subcutaneous fluid collections have been the predominant CT findings in cases of DSWI [47], but these features appear to be more specific and sensitive for DSWI presenting more than 3 weeks after surgery [48].

7. Morbidity and mortality

Mediastinal infection negatively impacts early, mid-term, and long-term survival after adult heart surgery [2, 7-12]. While it is intuitive that clinically serious DSWI reduces 30-day and/or in-hospital survival relative to similar patients not suffering this complications, the effect of DSWI on long-term survival is especially insulting since many patients referred for heart surgery expect to gain a survival advantage compared with other treatment options for their underlying heart disease [49].

Prior to the development of modern protocols for DSWI management, which include thorough sternal debridement and use of vascularized flaps to repair the mediastinal defect resulting from debridement, early mortality from DSWI exceeded 50% [50]. In the “modern era,” reported rates of in-hospital or 30-day DSWI mortality for range from 7% to over 30% [11,14, 35, 42, 51, 52]. In our own recent experience, early mortality for DSWI is approximately 16% [14]. Therefore, despite many advancements in intensive care medicine, DSWI continues to be a deadly complication. Death in the early period is typically the result of sepsis or other infectious complications including multiorgan failure [12]. Morisaki et al. recently demonstrated methicillin-resistant S. aureus infection to be an independent risk factor for in-hospital mortality in their cohort of poststernotomy DSWI patients [53].

Patients surviving infectious complications and the acute insult of DSWI exhibit reduced mid-term survival compared with controls. For example, one-year mortality for DSWI following CABG is significantly increased compared with similar patients who did not develop DSWI [2, 8, 10]. Milano et al and Braxton et al both demonstrated a doubling of mid-term mortality among patients with DSWI after CABG compared with controls [9, 54]. Karra et al examined predictors of one-year mortality after treatment for DSWI and found that delay in closing the mediastinal defect; age over 65 years; need for ICU care prior to sternal debridement; and methicillin-resistant S. aureus infections were each independently associated with mortality [7].

Long-term survival has consistently been demonstrated to suffer in patients with a history of poststernotomy DSWI [5, 8, 9, 12, 35, 52, 54-56, Figure 1]. For example, Filsoufi et al reviewed nearly 5,800 adult heart surgery patients at a single institution over 8 years and found DSWI to be associated with significantly reduced 5-year survival compared with patients who did not develop DSWI [35]. Similarly, Risnes et al reviewed their experience of over 18,000 cardiac surgical patients with a mean follow-up of over 10 years. Long-term survival for patients whose course was complicated by DSWI was <50% compared with >70% for patients without DSWI, and DSWI was independently associated with reduced long-term survival after cardiac surgery [HR 1.59; 95% CI 1.16 - 2.70, p = 0.003] [12]. Similar data have been also been reported by Toumpoulis et al. [55].
It is unclear as to the specific reasons for worse long-term survival after DSWI is successfully treated, but cardiac-related deaths or progression of cardiovascular disease appears to be a common cause of death for those with DSWI [8, 12]. In addition, Chu et al recently reported that patients with peripheral vascular disease had worse outcomes long-term after coronary artery surgery [57]. We recently evaluated long-term survival in a cohort of 222 adult cardiac surgical patients treated at Duke University Hospital for poststernotomy mediastinitis. Using multivariable regression analysis and at a mean follow-up of 5.5 years, the following variables were noted to be independently associated with survival: heart failure [Hazard ratio (HR) 1.58, p = 0.029]; sepsis [HR 2.38, p < 0.001]; peripheral vascular disease [HR 2.06, p = 0.001], age > 65 years [HR 1.61, p = 0.037]; and take back for bleeding [HR 2.96; p = 0.007] (unpublished data).

Not only does DSWI lead to increased mortality among post-surgical patients, it is also closely associated with other types of postoperative complications. For example, in a recent review of morbidity following coronary surgery from the Virginia Cardiac Surgery Quality Initiative, DSWI was the not only the most expensive complication, but it was also associated with a longer length of hospital stay, by more than two full weeks, than any other single postoperative complication, including perioperative stroke and renal failure [4]. Other reports have corroborated the impact of DSWI on overall hospital length of stay [2, 6, 52]. Speir et al also demonstrated that DSWI was the most likely form of postoperative morbidity to be associated with other complications such as prolonged ventilation, bleeding, renal failure, and atrial fibrillation [4]. DSWI has also been associated with increased rates of stroke, need for inotropic or mechanical cardiac support, and perioperative myocardial infarction, renal failure, and prolonged mechanical ventilation [12, 51].

8. Microbiology

Numerous bacterial pathogens may contribute to DSWI, but the most commonly isolated organisms are *Staphylococcus*, either coagulase-negative *Staphylococcus* (CNS) or *S. aureus*
In our own recently described experience, methicillin-sensitive and methicillin-resistant \textit{S. aureus} each accounted for 35\% of wound isolates, while CNS was present in 12\%, and gram negative organisms were found in 18\% [14]. It is noteworthy that our protocols for preoperative preparation of the cardiac surgical patient do not routinely incorporate mupirocin treatment. Additionally, many patients included in our cohort of patients treated for DSWI underwent primary cardiac surgery at referring facilities. Therefore, specific details in these patients predisposing to a certain pathogen were often lacking. Risk factors for the development of methicillin-resistant \textit{S. aureus} poststernotomy mediastinitis include previous hospitalization and takeback for bleeding after the original cardiac surgical procedure [59]. Gardlund et al have attributed the development of \textit{S. aureus} infection to the development of bacteremia [60], which is corroborated by our own data in which 50\% of patients treated for DSWI had evidence of bacteremia or septicemia (Jones classification 3b) at the time of clinical presentation with DSWI [14]. Finally, it has been demonstrated that methicillin-resistant strains of \textit{S. aureus} are more virulent and deadly than methicillin-sensitive strains, and, in fact, methicillin resistance was the only independent predictor of early mortality in one analysis [61].

In contrast, \textit{S. epidermidis} often presents with a rather indolent course, often not manifesting clinically until 3 weeks or more postoperatively [62]. CNS is frequently isolated in cases of sternal instability or nonunion, which may itself arise from obesity or acute or chronic pulmonary disease [60]. CNS has recently been noted to of increasing incidence and has been reported to be present in roughly 50\% of sternal infections in some series [6, 11, 60].

Gram negative organisms contribute less commonly to the pathogenesis of DSWI. Gram-negative infections of the mediastinum classically arise in the setting of other postoperative infectious complications such pneumonia, urinary tract infections, intra-abdominal infections, or other nosocomial infections [58, 60]. Mekontso-Dessap et al reviewed their experience with DSWI after cardiac surgery and found that Enterococcus species were the most common isolates in early cases of DSWI (<14 days), which likely arise from translocation of these bacteria from other sources in the host [63]. In contrast, \textit{Staphylococcus} species, often CNS, were the most common isolates in cases presenting more remotely from the original surgery [63].

\section*{9. Cost of DSWI}

One of the most impactful consequences of DSWI is the costs incurred in its management, in part due to the multiple associated conditions that frequently complicate the clinical picture of these sick patients [4, 12]. The average hospital costs for patients treated for DSWI is approximately 2.5 to 3 times that of similarly matched patients who enjoy an uncomplicated postoperative course [23, 64, 65]. As noted, increased costs are primarily attributed to associated comorbid conditions that arise during the treatment of mediastinitis, increased length of hospital stays, including frequent need for ICU services, and the need for multiple surgical procedures when using traditional approaches to DSWI [4, 23]. For example, Hollenbeak et al found that patients with DSWI remained hospitalized an average of 3 weeks longer than noninfected patients after coronary artery surgery [2]. Interestingly, in the same study it was noted that DSWI patients who ultimately expired incurred nearly $61,000 US dollars more in costs than other DSWI patients who ultimately survived, presumably secondary to the multiple comorbidities associated with clinically aggressive mediastinitis [2]. Using more recently acquired information, Speir et al found that mediastinitis was the single
most costly complication in Virginia after coronary artery surgery, raising the cost of care by over $62,000 US dollars, or more than 240% increase, on average, [4]. Therefore, it is apparent that treatment of these critically ill patients is extremely expensive to the hospital system, even if the complication is uncommonly encountered. As a result, and based on increased scrutiny of various hospital-associated conditions, the US Center for Medicare and Medicaid Services no longer reimburses for hospital costs incurred in treatment of DSWI following coronary artery bypass surgery [66]. Interestingly, however, it appears that use of negative pressure therapy in the treatment of DSWI contains associated costs. For example, Mokhtari et al reported the use of VAC therapy to be cost effective and efficacious in eradicating mediastinal infection [65]. In our own experience, patients treated with negative pressure therapy for DSWI had costs that were $150,000 US dollars less than the average Medicare charges for treatment required for DSWI ($152,00 vs. $ 300,000) [67].

10. Treatment of DSWI

Upon establishing the diagnosis of sternal wound infection, the immediate goal of treatment is complete eradication of infection followed by stabilization of the sternum and chest wall. Multiple strategies for managing mediastinitis have been proposed and range from open mediastinal packing to debridement with closure over drains, to placement of vascularized tissue flaps. In addition, negative pressure therapy (NPT) has been used to aid in the treatment of mediastinal infection, as first reported in 1999 [68]. Several initial series described the use of NPT for treating mediastinitis and outlined potential benefits of this approach to controlling mediastinal infection [15, 16, 45, 69, 70]. However, a consensus as to best treatment for mediastinal infection has yet to be established. For example, Schimmer et al recently surveyed 79 cardiac surgery programs in Germany to query DSWI management strategies [71]. They found that approximately 1/3 of the centers preferentially used NPT for controlling mediastinal contamination, while another 1/3 use closed chest irrigation, and another 1/3 combine management approaches [71]. Other groups continue to advocate more aggressive surgical therapy in addressing infected sternal wounds [7, 72].

Initially, the primary treatment for DSWI was open packing followed by secondary closure [73]. This method was labor intensive, cumbersome, and was characterized by significant rates of recurrent infection and other complications [58]. Shortly thereafter, Schumaker and Mandelbaum proposed early closure over drains [74], which allowed continual irrigation of the infected mediastinum [73, 74]. Although this approach represented an improvement relative to open packing procedure, the latter procedure was still associated with failure rates of at least 25% [73].

Omental flap repair of the infected mediastinum was originally reported in 1976 by Lee [75] and has been shown to have distinct advantages over continuous antibiotic irrigation through drains [76] and over muscle flaps for sternal wound defect repair [9, 14]. This is thought to occur by obliterating dead space, a mechanism that can also be credited to muscle flap repair of the open mediastinum [26]. However, the omentum may have angiogenic and immunologic properties that stimulate more complete sternal defect healing [58]. Other advantages of using the greater omentum to flap the sternal defect include superior malleability and excellent blood flow. As originally described, traditional omental flap repair of the mediastinum involves a laparotomy to mobilize and place the graft through the anterior portion of the diaphragm [Figure 2]. More recently, harvesting the
Fig. 2. Preparation of the omental flap graft for transfer to the mediastinum from Milano et al with permission [80]. After preparation of the mediastinum [A], a laparotomy is performed and the greater omental graft is prepared by mobilizing it from the transverse colon [B]. Once the graft is tunneled through an anterior diaphragm incision, it is secured in the mediastinum to fill the defect. Care is taken to avoid twisting or kinking the omental graft as it is delivered from the abdomen into the mediastinum [C]. The mediastinum is then closed over drains with retention sutures. The drains are placed to bulb suction [D].

Omental pedicle using a laparoscopic approach may have advantages over open abdominal procedures and has been utilized successfully [77]. Jurkiewicz et al first reported on the successful use of muscle flaps in DSWI in 1980 [78], while the Emory groups’ experience with muscle flap repair of the infected mediastinum refined approaches and clarified advantages of these techniques [26]. As with omental flap repair, discrete advantages of complete mediastinal repair with vascularized muscle flaps were demonstrated as compared with closed chest irrigation [26, 79]. Milano et al compared omental and pectoralis muscle flap repairs for the infected mediastinum and found that procedural details, early complications, and hospital length of stay were all improved with omental flap repair [80]. Omental flap repair also trended towards improved survival and reduced chronic pain when compared with pectoralis flap procedures [80]. In fact, significant long-term complications can be associated with muscle flap repair of the mediastinum including paresthesias, sternal instability, truncal weakness, and prolonged pain syndromes [9, 26, 81, 82]. Reporting on the long term results of muscle flap coverage of the mediastinum, Ringelman noted persistent pain in over 50% of patients, numbness and paresthesias in 44%, sternal instability in over 40%, and shoulder weakness in 1/3 of patients [82]. It is worth noting, however, that unfavorable outcomes with open packing and/or mediastinal irrigation prompted recommendations for aggressive mediastinal debridement including radical sternectomy.
[78, 83]. While effective in removing devitalized and infected tissue, the resulting sternal defect required large-volume vascularized tissue such as muscle flaps or large omental flaps to obliterate deadspace. As a result, many of the negative long-term consequences often experienced and attributed to muscle flap repair may actually have originated with radical sternal debridement. After the introduction of NPT management of mediastinal sepsis/infection, radical sternal debridement has been de-emphasized [14, 16, 83, 84]. Stated differently, the use of NPT to treat the infected mediastinum helps to avoid radical sternal debridement and likely avoids chronic syndromes previously seen and perhaps erroneously attributed to treatment methods.

Obdeijn et al introduced the use of negative pressure therapy (NPT) for treatment of the infected mediastinum after median sternotomy [68]. The introduction of vacuum-assisted closure (VAC®) technology [Kinetic Concepts Inc. USA San Antonio, TX], based on the application of negative or subatmospheric pressure to the wound, has improved management of DSWI, as demonstrated by several groups including our own, and is now considered a cornerstone in the management of these complex clinical scenarios [14, 15, 70, 73]. Negative pressure therapy appears to induce effective proliferation of the effectors of wound healing [85] removes wound exudates, improves regional blood flow [86], and reduces accumulation of inflammatory mediators such that earlier and more complete wound healing results [70]. Laboratory studies also show that NPT induces early wound healing through microdeformations within the wound, stimulating cell division, proliferation, and angiogenesis [73, 87]. One distinct advantage of the mechanisms of NPT is more thorough eradication of infection, stimulation of vigorous wound granulation, and the subsequent promotion of safe and effective sternal closure either primarily or with rigid osteosynthesis [11, 14, 88]. Wound treatment with NPT appears to be significantly lower rates of recurrent wound complications such as reinfection, seromas, or hematomas [14, 89, 90]. However, there is a recognized tendency for recurrent infection when MRSA is the inciting organism or with prolonged mechanical ventilation [91].

Clinically, NPT has been associated with significantly lower mortality rates in the acute management of mediastinitis [11, 89]. For instance, Petzina et al recently compared 118 patients with poststernotomy DSWI and demonstrated that patients treated with NPT had better survival and less sternal re-infection compared with patients in whom NPT was not used [89]. In addition, as may have been expected, shorter hospital stays were noted within the NPT group. Baillot et al reported similar results by, while documenting a reduction in acute DSWI mortality from 14.1% to less than 5% when NPT was incorporated into treatment regimens [11]. Finally, De Feo et al evaluated 157 patients with poststernotomy DSWI at a single institution over 15 years. Patients in whom NPT was incorporated in the treatment regimen had reduced early mortality rates and, reduced reinfection rates, and slightly reduced overall hospital stays [92].

As noted previously, long term mortality among patients with DSWI has been historically poor compared with similar patients not suffering the complication, but recent evidence suggests this may be changing. The first evidence for this phenomenon was reported by Sjogren et al, who compared 46 patients with poststernotomy mediastinitis managed with NPT with a matched cohort of cardiac surgical patients not experiencing postoperative mediastinitis [93]. Actuarial and adjusted 5-year survival was not different between groups, demonstrating for the first time that long-term results of heart surgery were not negatively impacted by DSWI [Figure 3].
Fig. 3. Actuarial (panel A) and adjusted (panel B) survival curves for patients with poststernotomy mediastinitis treated with vacuum-assisted closure (VAC) as compared to control subjects who did not experience mediastinitis after heart surgery. No difference in long-term survival was demonstrated between the groups. From Sjogren et al [93], with permission.

Another study from the UK demonstrated that midterm survival for patients with postoperative mediastinitis was similar to patients not suffering the complication [51]. Similar results have been reported by Cayci et al from Columbia University, who found that DSWI was associated with increased early mortality but long-term survival was not different from controls. Furthermore, DSWI was not an independent predictor of mortality in their single-center experience [52]. It is unclear as to why this may be the case, but it is notable that NPT was used in over 80% of patients with mediastinitis as a
means for clearing infection, and nearly 50% of patients in their cohort were managed with vac therapy alone or vac + secondary sternal closure. In our own series, we found that use of NPT for controlling mediastinal infection was an independent predictor of survival on multivariable analysis [unpublished data], and that patients managed with NPT had significantly improved long term survival compared with patients not treated with NPT [Figure 4].

11. Suggested treatment algorithms for DSWI

While accumulating extensive experience with DSWI management as a referral center for these difficult problems. We and others have formalized protocols for managing mediastinal infection utilizing NPT which allows sternal salvage and improved outcomes in the majority of cases of DSWI [14-16, Figure 5].

![Kaplan Meier survival estimates](image)

**Log-rank p=0.008**

Fig. 4. Kaplan Meier survival curves for patients with poststernotomy mediastinitis treated with negative pressure therapy (NPT) or by traditional means for controlling mediastinal infection (controls). Patients treated with NPT had significantly improved long-term survival by log rank analysis.
Fig. 5. Suggested algorithm for management of mediastinal infection after cardiac surgery via median sternotomy, with permission from Sjogren et al [15]. Emphasis is placed on gentle sternal debridement, negative pressure therapy to the mediastinum, and closure based on monitored serum C-reactive protein levels.

As noted previously, successful treatment of DSWI begins with recognition of signs and symptoms of sternal infection, which may occasionally be subtle. Sternal wound exploration to ensure a prompt and accurate diagnosis is warranted when signs of mediastinitis are present. This approach also helps to distinguish between superficial and deep sternal infections. At the time of initial sternal exploration, tissues and fluid should be obtained for bacteriologic analysis. Targeted antibiotic therapy for 4-6 weeks duration is prescribed and is determined by the culture results [14, 15]. If the infectious process extends beneath the facial layer, all sternal hardware should be removed. The sternum itself is then gently debrided of grossly devitalized tissue, but wide excision of the sternum is not necessary and may be injurious and counterproductive. Limited sternal debridement is now preferred and good results have been seen with this approach [14, 16, 83, 93, 94]. Negative pressure therapy is then instituted on the opened incision after limited sternal debridement. The polyurethane foam is subsequently changed in the operating room or at the bedside every 2-3 days. During this time, assessment as to the state of the sternum is made to determine if mediastinal flap repair is required or if secondary sternal closure is possible. Our general approach for determining when the sternum can reliably be reapproximated is based on the state of the sternum after several days of NPT. Wound characteristics precluding secondary
sternal reapproximation include multiple transverse sternal fractures, poor bone stock, costosternal separation, or the requirement for such extensive sternal debridement that reapproximation of the sternal halves is not feasible. Gustafsson et al have advocated use of serum C-reactive protein levels to guide wound closure timing [95, Figure 6]. C-reactive protein levels less than 70 mg/L corresponded with successful sternal reapproximation [95]. Successes have also been reported with sternal plating as a treatment for the fractured sternum [11, 96], but our approach has been conservative in this regard since any residual infectious process could contaminate the implanted hardware [91]. Numerous clinical advantages for DSWI management protocols incorporating NPT have been observed, many centered upon the sternal stabilization achieved when vacuum-assisted closure is engaged [70]. The sternal stability afforded by NPT improves pain compared with open packing or other approaches for addressing the infected mediastinum [88]. In addition, the stabilized sternum yields several pulmonary benefits, the first of which is the ability to successfully separate from mechanical ventilation. This promotes earlier and more effective patient mobilization and prevents the patient from being confined to bed, where other complications common to DSWI therapy are often incurred [70]. Negative pressure therapy also improves ventilation and overall pulmonary function and leads to more effective chest physiotherapy [70, 97]. Importantly, no deleterious hemodynamic effects of NPT have been documented although this has been speculated [70].

12. Complications of treating mediastinitis

It has been estimated that approximately 15% of patients develop recurrent infection [98]. In the experience of Bapat et al, this has included recurrent infection with the same organism associated with the original sternal infection [99], and we speculate that if the polyurethane foam required for NPT is not adequately inserted with each dressing change, small, isolated spaces may arise within the wound that can become superinfected. We recently reported rates of recurrent wound complications associated with various mediastinal flap coverages. For example, muscle flap repair of the treated mediastinum, consisting predominantly of pectoralis muscle flaps, was associated with increased rates of recurrent wound complications such as hematoma, seromas, and recurrent infection [14]. Conversely, use of NPT prior to definitive repair of the sternal wound defect was associated with increased rates of successful secondary sternal closure without the need for any flap transfer, and with shortened length of hospital stay after definitive repair. Excellent results have been reported elsewhere when NPT is incorporated into management protocols for DSWI [65]. Petzina et al recently reported a 7.2% rate of “major complications” associated with NPT for DSWI in a cohort of 69 patients. Most complications were bleeding-related [100]. On the other hand, cardiac function and hemodynamics appear to be stable during NPT to the open mediastinal wound [101, 102].

When NPT is used, caution should be exercised with regard to the length of therapy. In our own experience, prolonged use of NPT leads to a “frozen” mediastinum, making subsequent closure by vascularized flaps or other technique difficult to perform and places the cardiac structures at risk for injury during subsequent sternal repair. Others have noted the similar difficulties [99]. In such cases, continued application of NPT to closure by secondary intent may be the best therapeutic option rather than to place the mediastinal structures at risk for injury during attempted flap repair.
13. Can mediastinitis be prevented?

Loop has stated: “prevention and better treatment of sternal wound complications must be a major goal in assuring the highest quality of cardiovascular care...[5]. Although most efforts towards DSWI have focused on the treatment of DSWI, several methods to reduce rates of mediastinitis have been proposed and validated recently. As a result, efforts to prevent mediastinal infection may already be working. For instance, investigators from Boston recently reported on their experience with DSWI between 1992 and 2006, separating analysis into early and late time periods. They noted that DSWI had decreased from 1.57% to 0.88% over the last 5 years of their analysis and attributed the positive findings to adoption of strict glucose control algorithms [40]. Tight glycemic control appears to be effective in significantly reducing rates of DSWI. [30, 103]. In addition, Lazar et al demonstrated improved coronary surgery outcomes with a strategy for strict glucose control (125 – 200 mg/dL) using glucose-insulin-potassium solution, including reduced ischemic events and improved rates of wound infection in a cohort of diabetic patients undergoing CABG [31].

Antimicrobial therapy has also positively impacted rates of DSWI. Most notably, appropriate timing and selection of preoperative antibiotics has been associated with reduced rates of surgical site infection [6, 104]. Furthermore, use of nasal mupirocin in patients undergoing cardiac surgery via median sternotomy eradicates 95 – 100% of S. aureus for up to one year postoperatively [105], and sternal wound infections are also reduced by nearly 2/3 in some series with the use of mupirocin in patients colonized with S. aureus [106, 107].

Technical details of the median sternotomy incision and closure almost certainly impact the likelihood for DSWI postoperatively. Baskett et al have argued that assiduously following technical details and proper surgical and aseptic techniques can also dramatically reduce rates of poststernotomy infection [42]. They emphasize the importance of accurate reapproximation of the sternal halves and caution against the use of bone wax to gain sternal hemostasis [42]. Since sternal instability is often evoked as one mechanism contributing to the development of sternal infection, evaluation of the most effective sternal closure methods has been undertaken. For example, Schimmer et al compared standard closure techniques by transsternal or peristernal wiring with techniques using additional lateral wire reinforcement in the method described by Robicsek in a cohort of 815 high-risk patients [108]. There were no differences observed in the rates of sternal dehiscence or superficial or deep sternal wound infections, but they did show that more sternal wires placed for closure was associated with significantly reduced rates of DSWI [108]. Others have emphasized that rigid sternal closure techniques are preferred, particularly in those considered at high risk for sternal wound complications such as dehiscence and infection. These techniques are widely used in most surgical practices that incorporate osteotomy incisions. In fact, cardiac surgery is now the only discipline not routinely repairing osteotomies with rigid plating techniques [109]. Lee et al recently reported their experience with titanium plate fixation of the sternum in 750 patients at high risk for sternal wound complications, noting 97.6% freedom from sternal infection or dehiscence [110]. Levin et al have also introduced another form of rigid sternal closure as an alternative to wire circlage and highlighted advantages to this approach [111]. Although rigid sternal closure techniques have not been compared prospectively with wire circlage, cadaveric studies have shown rigid plate fixation techniques to be superior to wire circlage by providing increased stiffness to the wound closure and less lateral displacement of the sternal halves [112].
Other suggested techniques for reducing the incidence of DSWI include ensuring true midline sternotomy as this is thought to preserve periosteal blood flow and limits transverse sternal fractures which contribute to sternal nonunion and instability [42, 44]. Since sternal ischemia is thought to play an important role in most cases of DSWI, limiting the length of internal mammary artery harvested for use as a coronary artery bypass conduit or avoiding its use altogether in prohibitively high risk patients may help to reduce DSWI [44]. Harvesting the internal mammary artery as a skeletonized conduit is also preferable to harvesting the graft as a pedicle since this preserves more peristernal blood flow [113]. Finally, sternal foreign bodies that may impede bony union, especially bone wax, should be avoided [42, 44]. In contrast, vancomycin paste or gentamicin-soaked absorbable sponges applied to the sternum upon closure have been shown to reduce rates of sternal bacterial contamination [114, 115]. However a more recent randomized controlled trial evaluating the impact of gentamicin-soaked collagen sponge on poststernotomy wound infection did not show an advantage of this approach over controls [116].

Based on the aforementioned successes with NPT in the treatment of documented DSWI we and others have recently evaluated the application of NPT to clean, closed incisions as a method to prevent complications in high-risk wounds [117, 118]. We initially applied this form of “well wound therapy” in a cohort of 57 adult cardiac surgery patients known to be at increased risk for DSWI based on a validated risk stratification model [13]. No cases of superficial or deep sternal wound infections were noted although the group had an estimated 6% risk for DSWI [118]. This form of NPT was noted to be easy to apply, well-tolerated by the patients, and was also judged to be cost-effective when utilized in patients with increased demonstrated risk for DSWI [117]. Since the time of the original report, we have used this novel wound treatment system in over 200 high risk patients and continue to observe reduce rates of sternal wound complications (unpublished data). However, residual problems have been encountered with gross technical errors including off-center sternal incisions, sternal fractures, and costo-sternal separation.

The mechanisms underlying such positive clinical findings are not well understood, but based on well described mechanisms of vacuum-assisted therapy in open incisions, it is hypothesized that applying NPT to the closed incision also favorably affects wound perfusion. Therefore, we assessed peristernal perfusion after median sternotomy and under various degrees of reduced native sternal perfusion as a result of mammary artery harvesting using laser Doppler flowmetry, demonstrating that after median sternotomy and IMA harvesting, peristernal perfusion is significantly reduced and recovers little in the first 4 postoperative days [119]. However, NPT applied to the closed incision increases peristernal perfusion compared with controls regardless of the status of the ipsilateral IMA, providing a rare piece of physiologic evidence for the efficacy of NPT and supports use of NPT as a form of “well wound therapy,” particularly in patients at high-risk for sternotomy complications [119] These findings are clinically important and relevant, implying that NPT can augment peristernal soft tissue perfusion made relatively ischemic by IMA harvesting. Although this study does not address bony perfusion, per se, Fokin et al proposed that substrate diffusion through peristernal tissues may be an important mechanism to maintain perfusion in sternal wounds rendered ischemic by mammary artery harvesting until collateral blood supply to the sternum is well-established [120, 121]. If so, NPT may indeed augment sternal and/or periosteal perfusion via improved peristernal “diffusion” through improved soft tissue perfusion.
14. Conclusions

Deep sternal wound infections remain dreaded and deadly complications associated with cardiac surgery. However, incremental improvements have been made recently with regard to lowering rates of observed infections due to a variety of measures. In addition, it appears that management of DSWI with negative pressure therapy alleviates the impact of this condition on short- and long-term survival. Further investigation is needed to determine the potential impact of negative pressure therapy on closed incisions as a novel method to prevent sternal wound complications.

15. References


[66] Center for Medicare and Medicaid Services; http://www.cms.hhs.gov/HospitalAcqCond/


This book considers mainly the current perioperative care, as well as progresses in new cardiac surgery technologies. Perioperative strategies and new technologies in the field of cardiac surgery will continue to contribute to improvements in postoperative outcomes and enable the cardiac surgical society to optimize surgical procedures. This book should prove to be a useful reference for trainees, senior surgeons and nurses in cardiac surgery, as well as anesthesiologists, perfusionists, and all the related health care workers who are involved in taking care of patients with heart disease which require surgical therapy. I hope these internationally cumulative and diligent efforts will provide patients undergoing cardiac surgery with meticulous perioperative care methods.

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