

Descending necrotizing mediastinitis: A 10-year surgical experience in a single institution

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Objective: Early diagnosis and aggressive surgical drainage are very important for successful treatment of descending necrotizing mediastinitis. However, the surgical techniques used for this condition remain controversial. We report our 10-year experience of managing this devastating disease, focusing on the multidisciplinary, minimally invasive operative procedures and the unique bacteriologic factors in Taiwan.

Methods: Between January 1997 and January 2007, we retrospectively reviewed 18 patients with descending necrotizing mediastinitis who were treated in the National Taiwan University Hospital. Diagnosis and Endo classification were confirmed by computed tomography of the neck and chest.

Results: Eight women and 10 men were included in this study. The mean age was 57.8 ± 15.2 years. Cervical drainage was performed in the involved area in all patients. The methods for mediastinal drainage included transcervical ($n = 10$), video-assisted thoracic surgical drainage ($n = 6$), subxiphoid drainage ($n = 1$), and mediastinoscopy-assisted drainage ($n = 1$). We could not rescue 3 patients because of uncontrolled sepsis before surgery, for a mortality rate of 16.7%. *Klebsiella pneumoniae* uniquely represents the most common pathogen in diabetic patients ($P = .01$), leading to more complicated courses in older patients ($P = .04$) and requiring more surgical interventions ($P = .05$) than other pathogens.

Conclusion: Transcervical mediastinal drainage is first justified in patients with limited disease in the upper mediastinum. For those with involvement of the lower anterior mediastinum, an additional subxiphoid approach is suggested. Cervicotomy with video-assisted mediastinal drainage is an excellent combination for involvement of the posterior mediastinum and pleural space. *Klebsiella pneumoniae* uniquely represents the most important and threatening causative pathogen for diabetic patients with descending necrotizing mediastinitis.

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Acute mediastinitis is a serious infection involving the connective tissue that fills the interpleural spaces and surrounds the median thoracic organs. The most common causes of mediastinitis are esophageal perforations and infection after operations through sternotomy incisions. One of the most dreaded and lethal forms of mediastinitis is descending necrotizing mediastinitis (DNM), which is referred to as a diffuse necrotizing variety that occurs as a complication of odontogenic or cervicofascial infections spreading along the deep fascial planes into the mediastinum.^{1,2} As infection spreads along these planes, widespread cellulitis, necrosis, abscess formation, and sepsis may occur through a fulminant course. Delay of diagnosis and delayed or incomplete drainage of the mediastinum are the main causes for the high mortality rate in this life-threatening condition.³⁻¹⁴ Even with the use of computed tomographic (CT) scanning, aggressive drainage, and modern antibiotic treatment, the mortality rate of DNM remains high.^{2,4,14} Surgical management, particularly the optimal form of mediastinal drainage, remains controversial. Many investigators have recommended the advantages of invasive procedures, including median sternotomy, clamshell incisions, and routine thoracotomy.^{4,9,14-16} However, these

Abbreviations and Acronyms

- CT = computed tomography
- DNM = descending necrotizing mediastinitis
- VATS = video-assisted thoracic surgery

invasive methods may lead to unexpected results, such as massive tissue injury, osteomyelitis, dehiscence of the sternum, and other complications.^{12,15,16}

We report our 10-year surgical experience with 18 patients affected by DNM, focusing on the application of multidisciplinary, minimally invasive procedures according to the extent of disease. The unique bacteriologic characteristics in Taiwanese patients with DNM are also discussed.

Patients and Methods

Between January 1997 and January 2007, we retrospectively reviewed the case histories of 18 patients with DNM who were hospitalized and treated in the Division of Thoracic Surgery, Department of Surgery and Traumatology, National Taiwan University Hospital. The medical charts, including personal demographics, image studies, operative records, hospital course, morbidity, and mortality were carefully reviewed. In all cases, the criteria of Estrera and colleagues² were fulfilled, including (1) clinical manifestation of severe oropharyngeal infection, (2) demonstration of characteristic radiographic features of mediastinitis, (3) documentation of necrotizing mediastinal infection at operation or postmortem examination, and (4) establishment of a relationship between oropharyngeal infection

and development of the necrotizing mediastinal process. In each case, this relationship was clearly established. Specifically excluded from this study were patients with mediastinitis resulting from an esophageal perforation, because the diagnosis, treatment, and outcomes in this subset of patients are fundamentally different and widely studied.

In all cases, diffuse painful neck swelling was clinically obvious, and respiratory insufficiency with sepsis ensued, suggestive of mediastinitis. All patients received empirical broad-spectrum intravenous antibiotics as soon as the diagnosis was suspected. Once the causative pathogen was cultured, the antibiotics were adjusted according to the sensitivity test. Diagnosis was confirmed by cervicothoracic CT on an emergency basis, which showed neck infection with soft tissue infiltration and edema of muscle tissues, as well as downward involvement of the mediastinum. The patients were evaluated according to Endo and colleagues' classification⁵ classification of the degree of diffusion of DNM based on CT findings: in type I, the infection is localized in the upper mediastinum above the carina; in type IIA, it extends to the lower anterior mediastinum; and in type IIB, it extends to the lower anterior and posterior portions of the mediastinum (Figure 1).

The surgical management consisted of cervical and mediastinal drainage. The neck was approached through an incision anterior to the sternocleidomastoid muscle in the involved side(s). The involved cervical spaces were opened, drained, and debrided of necrotic tissue. Several procedures are available for mediastinal drainage, including transcervical mediastinal drainage, mediastinoscopy-assisted drainage, video-assisted thoracic surgery (VATS) drainage, and subxiphoid drainage. The method selected depends on the Endo classification.⁵ Each procedure involved radical surgical debridement of the mediastinum and pleura with complete excision and

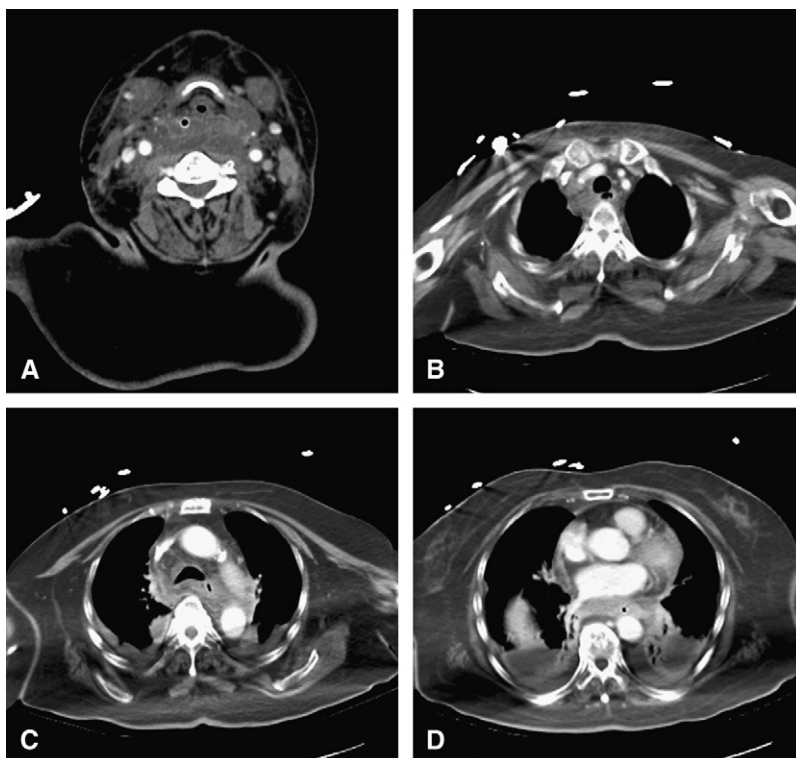


Figure 1. In case 16, the cervicothoracic CT scan revealed retropharyngeal and parapharyngeal abscess (A), descending into the upper mediastinum (B) and the posterior mediastinum with bilateral thoracic empyema (C and D).

decortication of necrotic tissue and adequate placement of silicone drains or chest tubes (Argyle 28 or 32; Sherwood Medical Company, St Louis, Mo) for drainage. The abscess and necrotic tissue from the deep neck area and mediastinum were sent for bacterial cultures intraoperatively. After primary surgery, follow-up CT scanning was performed in the surgical intensive care unit to assess the adequacy of therapy if clinical improvement was not obvious. Once residual abscess or necrotic tissue was detected, reoperation was immediately performed without hesitation. Repeated drainage was common owing to the progressive nature and great toxicity of DNM.^{6,9,11,12} The duration of drainage depended on clinical progress, return to normal CT scanning, and negative results of the cultures of fluids aspirated from the drainage tubes. We did not use routine irrigation of the mediastinum for fear of contamination and because there is not enough scientific evidence supporting this approach. We consider irrigation to be an optional method on the basis of qualitative evaluation of the material drained by the chest tubes.

Results

Our study comprises 8 women and 10 men. The characteristics, bacteriologic features, treatment course, and outcome of the patients with DNM are listed in Table 1. The mean age was 58.4 ± 15.1 years, ranging from 16 to 78 years. The foci of the inciting infections were retropharyngeal abscess in 5 (27.8%) patients, peritonsillar abscess in 4 (22.2%), odontogenic abscess in 3 (16.7%), acute epiglottitis in 3 (16.7%), foreign body in 2 (11.1%), and thermolysis catheter infection (Swan-Ganz; Edwards LifeSciences, Irvine, Calif) in 1 (5.6%). Fourteen (77.8%) patients had prior systemic diseases, including diabetes mellitus ($n = 11$, 61.1%), hypertension ($n = 8$, 44.4%), liver cirrhosis ($n = 2$, 11.1%), and coronary artery disease ($n = 2$, 11.1%). The delay between onset of the primary infection and operation varied from 1 to 8 days (mean, 2.7 days). Eleven (61.1%) patients were in Endo class I, 1 (5.6%) in Endo class IIA, and 6 (33.3%) in IIB.

Operative Management

All patients underwent cervicotomy for radical debridement. Among the 11 patients in Endo class I, 10 underwent transcervical mediastinal drainage and 1 received mediastinoscopy-assisted drainage. For the only patient in Endo class IIA, subxiphoid drainage was performed. All 6 patients in Endo class IIB underwent VATS mediastinal drainage. Seven (38.9%) patients underwent reoperation for postoperative CT showing residual abscess and necrotic tissue, including 2 patients undergoing a third operation. VATS reoperation was used in 4 patients, including 3 patients with type I disease who had primary transcervical drainage and 1 who had primary VATS drainage. Recervicotomy was performed in 5 patients, and tracheostomy was performed in 10 (55.6%) patients.

Clinical Outcome

The outcome was favorable in 15 (83.3%) patients, but 3 patients were not rescued, for a mortality rate of 16.7%.

The mean hospital stay was 40.1 ± 23.3 days (range, 9–90 days), and their mean intensive care unit stay was 14.6 ± 13.4 days (range, 1–41 days). Neither intraoperative death nor immediate postoperative death occurred. The 3 patients who died included 2 patients with significant comorbidity (both having diabetes mellitus and liver cirrhosis) and 1 with old age (>75 years old). Hospitalization time was marked in all 3 (4, 4, and 8 days), and sepsis was evident before the operation. Although immediate surgery was performed (including transcervical mediastinal drainage in 2 and VATS drainage in 1), sepsis with multiple organ failure still occurred.

Bacteriology

Bacteriologic results from intraoperative samples obtained from the neck and mediastinum showed positive culture results in 14 (77.8%) patients, including polymicrobial infection in 5 (35.7%) and monopathogen in 9 (64.3%) patients. The most common pathogen was *Klebsiella pneumoniae*, found in 6 (42.9%) patients, followed by viridans streptococci in 4 (28.6%) patients. Compared with non-*Klebsiella* DNM ($n = 8$), *Klebsiella* DNM ($n = 6$) occurred in significantly older patients ($P = .04$ by the Student *t* test) with a higher prevalence of diabetes mellitus ($P = .01$ by the Fisher exact test) and a higher rate of reoperation or mortality ($P = .05$ by the Fisher exact test) (Table 2).

Discussion

Nowadays, acute mediastinitis resulting from primary oropharyngeal or odontogenic infection is rare in Western countries. It is more common in developing countries owing to the poor economic conditions and consequent lack of medical resources for prevention and treatment of dental and oropharyngeal diseases. Reports indicate that this lethal complication will develop in about 2.6% of patients with deep neck infection in Taiwan.^{17,18} Our institution treated 18 patients in a period of 10 years, which indicated the incidence is not rare. During the past 50 years, efforts to reduce the mortality rate associated with DNM have been only moderately successful. In the first modern series of patients with DNM published in 1938, Pearse¹ reported that 49% of patients died during their treatment. However, despite the subsequent introduction of intravenous antibiotics, vast improvements in anesthesia and critical care, and the development of CT imaging, the frequency of death for patients with DNM reported in the literature over the past 3 decades has remained high in the antibiotic era.^{2,4,14}

Delay in diagnosis is the main reason for the high mortality rate in DNM because it usually runs a fulminant course.³⁻¹⁴ Contiguous cervicothoracic CT scan immediately confirms the diagnosis with high accuracy, showing soft tissue infiltration with loss of the normal fat planes or collection of fluid density with or without the presence of gas bubbles. It demonstrates the continuity of the infectious process between the

TABLE 1. Characteristics, treatment course, and outcome of the patients with DNM

No.	Age/sex	Systemic diseases	Etiology	Endo class	Bacteriology	Surgical treatment	Reoperation	ICU stay (d)	Outcome
1	34/M	None	Odontogenic abscess	I	Group F and viridans streptococci	B cervicotomy and transcervical mediastinal drainage	None	1	Discharge on POD 22
2	63/F	DM	Peritonsillar abscess	I	<i>Klebsiella pneumonia</i>	L cervicotomy and transcervical mediastinal drainage	None	15	Discharge on POD 63
3	48/M	None	Foreign body	I	Prevotella species, peptostreptococci	B cervicotomy and transcervical mediastinal drainage	None	4	Discharge on POD 36
4	50/M	None	Foreign body	I	<i>Streptococcus constellatus</i>	B cervicotomy and transcervical mediastinal drainage	None	3	Discharge on POD 39
5	56/M	HTN	Peritonsillar abscess	I	None	R cervicotomy and transcervical mediastinal drainage	None	1	Discharge on POD 28
6	65/M	DM	Peritonsillar abscess	I/A	<i>Klebsiella pneumonia</i>	L cervicotomy and subxiphoid drainage	Cervicotomy	3	Discharge on POD 15
7	57/F	DM, HTN, LC, PUD	Acute epiglottitis	I	<i>Klebsiella pneumonia</i>	L cervicotomy and transcervical mediastinal drainage	B VATS	22	Death on POD 22
8	65/F	DM, LC, HCC	Odontogenic abscess	I/B	<i>Klebsiella pneumonia</i>	L cervicotomy and B VATS	None	9	Death on POD 9
9	78/F	DM	Retropharyngeal abscess	I	Viridans streptococci	B cervicotomy and transcervical mediastinal drainage	2nd OP: B VATS 3rd OP: B cervicotomy and B thoracotomy	30	Death on POD 30
10	50/M	DM, HTN, CRI	Retropharyngeal abscess	I	β -Hemolytic streptococci	B cervicotomy and mediastinoscopic mediastinal debridement	None	5	Discharge on POD 23
11	16/M	None	Acute epiglottitis	I/B	Viridans streptococci	R cervicotomy and B VATS	None	6	Discharge on POD 30
12	78/F	DM, HTN, CAD, CVA, PUD	Retropharyngeal abscess	I	None	L cervicotomy and transcervical mediastinal drainage	None	13	Discharge on POD 32
13	52/M	None	Retropharyngeal abscess	I/B	β -Hemolytic streptococci and anaerobics	R cervicotomy and B VATS	B VATS	41	Discharge on POD 50
14	69/M	DM, HTN, COPD	Peritonsillar abscess	I	<i>Klebsiella pneumonia</i>	L cervicotomy and transcervical mediastinal drainage	L cervicotomy and R VATS	40	Discharge on POD 40
15	64/F	DM, HTN, CAD	Acute epiglottitis	I/B	None	R cervicotomy and R VATS	None	18	Discharge on POD 47
16	66/F	DM, HTN	Retropharyngeal abscess	I/B	None	B cervicotomy and B VATS	B cervicotomy	34	Discharge on POD 61
17	72/F	DM, HTN, SAH	Thermodilution catheter infection	I/B	<i>Klebsiella pneumonia</i>	R cervicotomy and B VATS	R cervicotomy for two times	10	Discharge on POD 90

Continued

TABLE 1. Continued

No.	Age/sex	Systemic diseases	Etiology	Endo class	Bacteriology	Surgical treatment	Reoperation	ICU stay (d)	Outcome
18	57/M	Oral cancer, panhypopituitarism with steroid usage	Odontogenic abscess	I	β -Hemolytic streptococci	L cervicotomy and transcervical mediastinal drainage	None	7	Discharge on POD 28

M, Male; F, female; DM, diabetes mellitus; HTN, hypertension; LC, liver cirrhosis; PUD, peptic ulcer disease; HCC, hepatic cell carcinoma; COPD, chronic obstructive pulmonary disease; CRI, chronic renal insufficiency; CAD, coronary artery disease; CVA, cerebrovascular accident; SAH, subarachnoid hemorrhage; VATS, video-assisted thoracoscopic surgery; R, right; L, left; B, bilateral; POD, postoperative day; OP, operation.

neck and thorax, establishing the relationship between neck infection and mediastinitis (Figure 1). Moreover, the CT scan demonstrates the extent of disease and stage evaluated by Endo's classification,⁵ which guided surgeons choosing the optimal operative procedures, ranging from transcervical drainage to posterolateral thoracotomy. Therefore, routine chest CT scanning was suggested for patients with deep neck infection.^{2,4-6,9,12,14,19-21} As in other types of necrotizing infections, intravenous broad-spectrum antibiotic therapy alone is not effective without adequate surgical drainage, extensive debridement, and excision of necrotic tissue.

Aggressive surgical drainage is essential for successful management of DNM, including cervical and mediastinal drainage. Cervical exploration and drainage, including definitive treatment of the inciting oropharyngeal or cervical infection, is considered compulsory therapy.^{2,4,6,9,14} However, because of the anatomic relationships of the cervicothoracic region, the progressive nature of DNM, and the variety of incisions available, controversy still exists as to whether a transthoracic approach for drainage and debridement is universally required and what the optimal surgical approach should be. Several approaches for mediastinal drainage have been proposed in the literature: transcervical, subxiphoid, median sternotomy, clamshell incision, posterolateral thoracotomy, video-assisted mediastinoscopy, and VATS.^{2,4,6-14,16,19,20,22-25}

Estrera and colleagues² described the first large series in the antibiotic era. On the basis of their own analysis, they supported transthoracic drainage for mediastinitis below the fourth thoracic vertebra posteriorly and below the carina anteriorly for persistent sepsis after cervical drainage. In

1990, Wheatley and colleagues¹⁴ condemned simple cervical drainage because their review of the literature revealed that transcervical mediastinal drainage was inadequate in 80% of patients. They preferred the combination of cervical drainage with anterior mediastinal drainage through the subxiphoid approach. Later, Marty-Ané,⁹ Corsten,⁴ Freeman,⁶ and their colleagues advocated aggressive mediastinal drainage via a standard thoracotomy approach as the optimum treatment for DNM, regardless of the level of mediastinal involvement. They stated that the mediastinum cannot be adequately drained via a limited approach through subxiphoid or anterior mediastinotomy and suggested the use of early thoracotomy for the best control of mediastinal sepsis. Moreover, some advocated the clamshell incision owing to excellent exposure for bilateral decortication and debridement of the entire mediastinum, including pericardiectomy.¹⁵ Although the exposure is excellent with the advantage of a 1-stage operation, it is a high-risk approach for critically ill patients with overwhelming sepsis and it exposes them to the risk of phrenic nerve palsy and sternal osteomyelitis.¹² Median sternotomy also seems feasible in type I and IIA DNM but inadequate in type IIB DNM, because the approach to the posterobasal compartments of the chest cavity is difficult.¹⁶ Subsequent osteomyelitis and dehiscence of the sternum were also dreaded complications. However, a vacuum-assisted closure device may be useful, because it promotes tissue approximation and stimulates the ingrowth of granulation tissue in sternotomy.²⁵

In 1999, Endo and colleagues⁵ classified DNM into types I, IIA, and IIB according to the degree of diffusion of the

TABLE 2. Comparison of patients with non-Klebsiella and Klebsiella DNM

	No. of patients	Mean age \pm SD (y)	No. of diabetic patients (%)	No. of patients with reoperation or mortality (%)	No. of patients with tracheostomy (%)	ICU stay \pm SD (d)	Hospitalization \pm SD (d)
Klebsiella DNM	6	65.2 \pm 5.2	6 (100)	5 (83)	4 (67)	16.5 \pm 13.2	40 \pm 31.3
Non-Klebsiella DNM	8	48.1 \pm 17.8	2 (25)	2 (25)	3 (38)	12.1 \pm 14.8	32 \pm 9.8
P values		.04*	.01†	.05†	.29†	.58*	.5*

SD, Standard deviation; ICU, intensive care unit. *Student *t* test. †Fisher exact test.

infection. They suggested that transthoracic mediastinal drainage might not always be required in type I but was always needed in type II. Because of the vigorous application of VATS in thoracic surgery worldwide since 1990, more and more authors advocate it as one of the treatments for this devastating disease. Roberts and colleagues²⁶ first reported a case of thoracoscopic drainage as an alternative way in a patient with mediastinal abscesses resulting from esophageal perforation. In 2004, Isowa and colleagues⁷ reported the successful management of a patient with DNM via VATS. Furthermore, two other groups of authors reported their successful use of VATS in 4 and 9 patients with DNM and emphasized the lower degree of invasiveness and early application.^{10,24} Because VATS allows excellent visualization of the entire thoracic cavity, it is possible to drain mediastinal, pleural, or pericardial effusions in the course of treating DNM (Figure 2). Sometimes, single-lung ventilation was difficult or impossible in critically ill patients with DNM. In that circumstance, we performed VATS debridement and drainage with double-lung ventilation. Alternatively, we induced short-term apnea or ventilation with a small tidal volume. The above strategies made the operation possible.

On the basis of our experience, the stage and the extent of DNM, as well as the clinical condition of the patients, should be carefully pondered when operative methods are being chosen. In our country, more than half of the patients have type I DNM, possibly because Taiwan is a small island and medical service is very convenient, which might be helpful in early diagnosis. When type I DNM is noted, cervicotomy with transcervical mediastinal drainage may be sufficient. Mediastinoscopy-assisted drainage may also be helpful in

this stage.¹³ Although most of the patients (8/11, 73%) with type I DNM were treated successfully by transcervical drainage without a transthoracic approach, 3 (27%) of them still needed VATS reoperation later. A low threshold should exist for performing a VATS procedure even in Endo type I cases when there is even a remote concern regarding caudal extension of the septic process to or below the carina. When type IIA was noted, an additional subxiphoid approach was suggested. When applied early in type IIB, VATS may be an excellent tool. Reoperation and bilateral involvement are likely owing to both the great toxicity of DNM and the weak immunologic status of patients. VATS can play an ideal role both for initial treatment and for reoperation because of its lower degree of invasiveness. Less invasiveness and surgical trauma bring less impairment of the immunologic status and a lower inflammatory and cytokine response,^{27,28} which are also beneficial given the critical condition of patients with DNM. In addition, VATS has the universal advantages of minimally invasive surgery, such as little pain, better cosmesis, and faster recovery.^{10,13} Our policy of multidisciplinary, minimally invasive procedures achieved a low mortality rate of 16.7%, which was encouraging.

The incidence of DNM is much higher in patients who are in poor physical condition owing to malnutrition, diabetes mellitus, alcoholism, drug addiction, or immunosuppression than in the general population.^{4,9,20,23} Diabetes mellitus is the most common systemic disease in DNM patients in our study, with an incidence of 61.1%. It has been reported that, in Taiwan, several infectious diseases in diabetic patients are most frequently caused by *Klebsiella pneumoniae*, including pyogenic liver abscess, empyema, meningitis, necrotizing fasciitis, and deep neck infection.^{18,29-35} As for

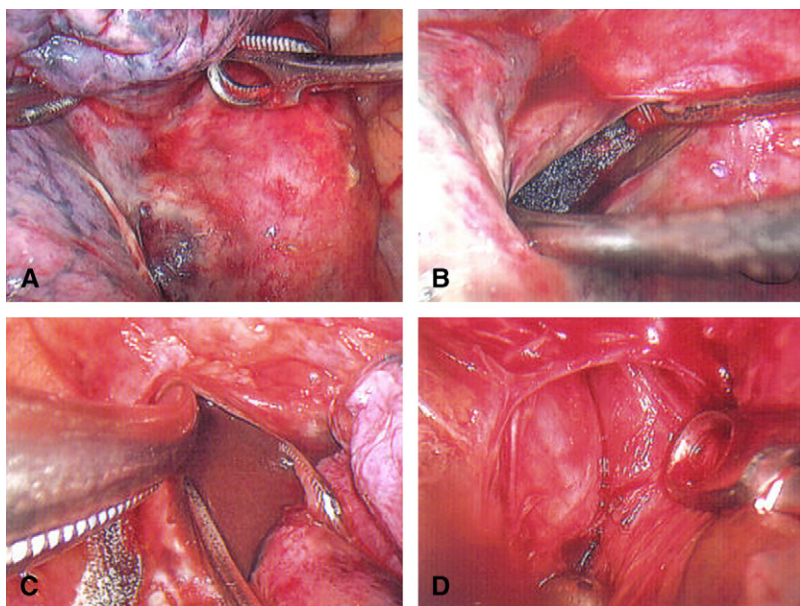


Figure 2. In case 16, during left VATS exploration, the inflamed mediastinal pleura at the para-aortic area was opened (A and B). Much purulent discharge and necrotic tissue was drained out and debrided with normal saline irrigation (C). Finally, the mediastinum was clear (D) and chest tube placement was precise with VATS.

DNM, *Klebsiella pneumoniae* was also the most frequently isolated sole pathogen in our study. Previous studies revealed that the bacteriologic pattern of DNM is usually polymicrobial and most commonly noted with beta-hemolytic streptococci,^{2,4,6} but our study showed only 35.7% were polymicrobial. Moreover, our finding demonstrates a significant correlation between diabetes mellitus and *Klebsiella pneumoniae* in patients with DNM ($P = .01$). The reason for the association of *Klebsiella pneumoniae* infection with diabetes mellitus remains unknown, but the infections caused by *Klebsiella pneumoniae* in diabetic patients are more severe and require more aggressive intervention and repetitive drainage.^{17,30,33} The *Klebsiella* DNM in our study showed the same tendency, being associated with older patients ($P = .04$) and a higher rate of reoperation or mortality ($P = .05$). Therefore, in clinical practice, we should be very alert about the diagnosis of DNM, especially in diabetic patients, because an even more fulminant course caused by *Klebsiella pneumoniae* might be encountered, which requires quicker and more aggressive surgical intervention.

Conclusions

DNM remains a life-threatening infection. On the basis of experience accrued in treating 18 patients in a period of 10 years, we conclude that a laddered approach with transcervical mediastinal drainage is first justified in patients with limited disease in the upper mediastinum, but a low threshold should exist for performing early VATS. For those cases involving the lower anterior mediastinum, an additional sub-xiphoid approach was suggested. Cervicotomy with VATS mediastinal drainage is an excellent combination for involvement of the posterior mediastinum and pleural space. *Klebsiella pneumoniae* uniquely represents the most common pathogen in diabetic Taiwanese patients, leading to more complicated courses in older patients and requiring more surgical interventions than other pathogens.

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