

A comparison between two lung ventilation with CO₂ artificial pneumothorax and one lung ventilation during thoracic phase of minimally invasive esophagectomy

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Background: To investigate the feasibility and safety of two lung ventilation with artificial pneumothorax in minimally invasive esophagectomy (MIE) through a comparison with conventional one lung ventilation.

Methods: Eleven hundred and sixty-six patients with esophageal cancer, who underwent McKeown MIE in our center from February 2006 to December 2016, were studied retrospectively. Seven hundred and five patients who underwent one lung ventilation with double lumen endotracheal tube (DLET) were assigned to DLET group. Other 461 patients who underwent two lung ventilation with single lumen endotracheal tube (SLET) were assigned to SLET group. Clinical characteristics, surgical variables and complications were compared between two groups.

Results: There were comparable patient characteristics in two groups. Surgical variables and complications were discussed between two groups. SLET group seemed to have shorter operative time, shorter postoperative hospital stay, and more harvested recurrent laryngeal nerve (RLN) lymph nodes than DLET group, which might be attributed to experienced surgeons. However, there were no significant differences of complications between two groups. Intraoperative clinical parameters were further studied. Before intubation and artificial pneumothorax, there were no significant differences between two groups, except diastolic blood pressure (DBP). With the application of artificial pneumothorax, patients in SLET group have obviously higher PO₂, PCO₂, and PetCO₂ value, and slightly lower pH value and blood pressure during thoracic phase. After the thoracic phase, the changes induced by artificial pneumothorax in SLET group were gradually reversed and clinical parameters gradually return to normal level.

Conclusions: Two lung ventilation with artificial pneumothorax is a safe and feasible choice during MIE.

Keywords: Two lung ventilation; single lumen endotracheal tube (SLET); minimally invasive esophagectomy (MIE); esophageal cancer; artificial pneumothorax

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Introduction

Esophageal cancer is one of the most common cancers in china, whose incidence rate and mortality rate were as high as 4.8‰ and 3.8‰ in 2015 (1). Surgical resection is considered to be the most effective treatment for early stage esophageal cancer. Researches indicated that open esophagectomy is related to significantly higher morbidity of perioperative complications, while minimally invasive esophagectomy (MIE) is proved to have better short term outcome and has gradually become a standard surgical approach for esophageal cancer in recent years (2,3).

Double lumen endotracheal tube (DLET) is commonly adopted for the anesthesia of MIE, which allows one lung ventilation in order to provide enough surgical space and good exposure. However, disadvantages of DLET anesthesia include rigorous requirement of pulmonary function, sophisticated procedure of intubation, high incidence of displacement during surgery, and frequent respiratory complications after surgery (4,5). Some patients with esophageal cancer are smokers and suffer from chronic respiratory diseases. These patients with poor pulmonary function have higher incidence of perioperative respiratory complications. Some even cannot bear one lung ventilation. Single lumen endotracheal tube (SLET) with CO₂ insufflation artificial pneumothorax, which is an alternative for DLET, allows two lung ventilation during surgery, and provides good exposure as well. However, Reports of SLET anesthesia in MIE are rare. Only Zhang *et al.* and Cai *et al.* reported 42 and 83 cases (6,7). In video-assisted thoracoscopic surgery (VATS), most application of two lung ventilation with SLET are found in endoscopic sympathectomy and spontaneous pneumothorax which do not require sophisticated surgical operation (4,8-13). As one of the largest esophageal center in china, we adopted SLET with CO₂ insufflation artificial pneumothorax and two lung ventilation in MIE since 2013. Here, we retrospectively studied patients who underwent MIE surgery in our center from February 2006 to December 2016, and assessed the safety and feasibility of application of SLET with CO₂ insufflation artificial pneumothorax in MIE.

Methods

Patients

Eleven hundred and sixty-six patients with esophageal cancer, who accepted minimally invasive McKeown three-phase esophagectomy in our center from February 2006 to

December 2016 (*Figure S1*), were retrospectively studied. Seven hundred and five patients of them, who accepted general anesthesia with one lung ventilation using a DLET during the thoracic phase of surgery, were assigned to DLET group. Other 461 patients accepted anesthesia with two lung ventilation using a SLET and CO₂ insufflation (CO₂ pressure =8 mmHg) artificial pneumothorax and were assigned to SLET group. This study has been approved by the Ethics Committee of Zhongshan Hospital (No. B2017-153), and all patients signed informed consent.

Anesthetic and surgical procedure

Combined general-epidural anesthesia is adopted. All the patients adopted the minimally invasive McKeown three-phase esophagectomy (14), which includes the extracorporeal gastric conduit creation and cervical anastomosis between gastric conduit and esophagus. Patients were placed in the left lateral prone position during the thoracic phase and in the supine position during other two phases. The changes of heart rate (HR), blood pressure, SpO₂, PetCO₂ were collected from the anesthesia note. During surgery, artery blood sample was taken for blood gas analysis once in the thoracic phase (generally 30–60 minutes after the initiation of thoracic phase) and again after thoracic phase (generally 30–60 minutes after thoracic phase) by anesthetist. These data were also collected from the anesthesia note.

Statistical analysis

Data are presented as the mean \pm SD. All statistical analysis was performed using SPSS 13.0, with $P < 0.05$ considered statistically significant.

Results

Clinical characteristics of two groups were summarized in *Table 1*. There were no significant differences in clinical characteristics between two groups, including age, gender, American Society of Anesthesiologists (ASA) grading, smoker number, TNM staging, tumor location, pulmonary function, and concomitant disease.

Surgical variables and postoperative complications were compared between two groups (*Table 2*). SLET group seemed to have shorter total operative time, shorter thoracic phase time, shorter postoperative hospital stay, more harvested left recurrent laryngeal nerve (RLN) lymph

Table 1 Characteristics of patients

Variables	SLET group (n=461)	DLET group (n=705)	P value
Age (years)	61.40±7.92	61.80±7.89	0.40
Gender			0.25
Male	359	527	
Female	102	178	
ASA			0.79
I	279	440	
II	147	216	
III	35	49	
IV	0	0	
Smokers	267	437	0.18
TNM stage			0.17
I	161	219	
II	193	334	
III	107	152	
Tumor location			0.97
Upper 1/3 thoracic oesophagus	53	84	
Middle 1/3 thoracic oesophagus	251	385	
Lower 1/3 thoracic oesophagus	157	236	
PFT's			
FEV1 (Act/Pred %)	84.47±17.76	84.64±18.40	0.88
DLCO (%)	81.28±16.80	80.50±17.43	0.45
Concomitant disease			
Cardiovascular disease	27	39	0.92
Respiratory disease	34	53	0.98
Diabetes	16	27	0.87

SLET, single lumen endotracheal tube; DLET, double lumen endotracheal tube; ASA, American Society of Anesthesiologists; DLCO, diffusing capacity of the lung for carbon monoxide; PFT, pulmonary function test.

nodes, and more total RLN lymph nodes than DLET group, which might be attributed to experienced surgeons in 2013–2016. However, there were no significant differences of complications between two groups. SLET group seemed to have the symptom of intraoperative hypotension during thoracic phase in a few patients, which could be relieved immediately by the removal of CO₂ insufflation.

Clinical parameters, including HR, systolic blood pressure (SBP), diastolic blood pressure (DBP), SpO₂, PetCO₂, and results of blood gas analysis (pH, PO₂, PCO₂)

were investigated in three time points of surgery (before anesthesia, during thoracic phase, and after thoracic phase). Before anesthesia, there were no differences of these parameters between two groups, except the DBP. During thoracic phase, patients in SLET group had lower blood pressure, and pH value, but higher SpO₂, PO₂, PCO₂, and PetCO₂ than patients in DLET group. After thoracic phase, most of these changes were alleviated. Most variables showed no significant differences between two groups. Significant differences were only observed in SpO₂ and pH value.

Table 2 Comparison of surgical variables and postoperative complications

Variables	SLET group (n=461)	DLET group (n=705)	P value
Total operative time (min)	191.12±29.02	214.01±35.23	<0.01
Thoracic phase time (min)	67.08±22.34	74.32±27.50	<0.01
Left RLN nodes (n)	3.92±1.99	2.40±1.76	<0.01
Total RLN nodes (n)	6.94±2.11	4.19±2.30	<0.01
Postoperative hospital stay (days)	10.21±9.03	11.41±9.41	0.03
Complications			
Anastomosis leak	53	81	0.93
Respiratory complications	26	43	0.84
Chylothorax	4	6	0.77
Hoarseness	24	36	0.95
Arrhythmia	19	28	0.98
Haemorrhage	3	6	0.97
Postoperative mortality	2	3	0.66
Intraoperative hypotension	3	0	0.12

SLET, single lumen endotracheal tube; DLET, double lumen endotracheal tube; RLN, recurrent laryngeal nerve.

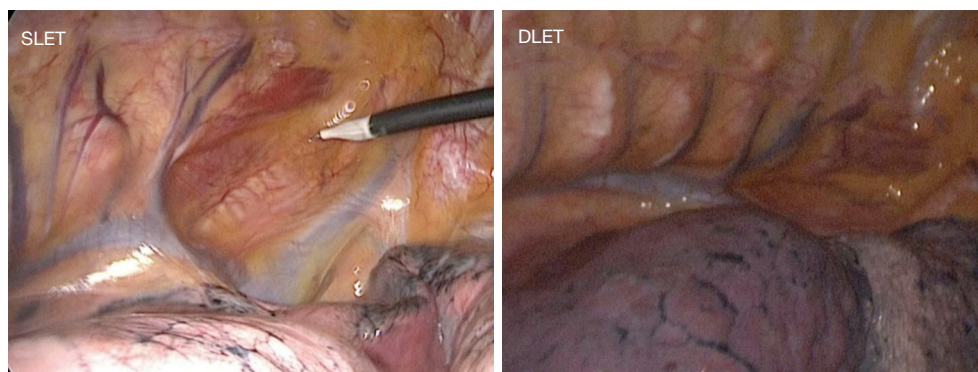


Figure 1 Good exposure with two lung ventilation. With the press on lung by the assistant, two lung ventilation with SLET could have good surgical exposure as one lung ventilation with DLET. SLET, single lumen endotracheal tube; DLET, double lumen endotracheal tube.

Discussion

Originally, one lung ventilation with DLET was the only choice during the anesthesia of patients who accepted MIE for the purpose of good exposure. However, researches have indicated some disadvantages of one lung ventilation with DLET (15-17), including airway injuries, intraoperative hypoxaemia, tracheobronchial injury, and so on. Many institutions have attempted to replace DLET with SLET in MIE nowadays, in order to avoid these defects. However,

it's still controversial whether patients benefit from SLET with artificial pneumothorax. Most studies were too small to provide meaningful statistical results.

One defect of one lung ventilation with DLET is complicated preoperative intubation and perioperative management. During the placement of DLET, the assistant of a fiber-optic bronchoscope is necessary. Besides, intraoperative translocation of DLET is frequent observed, and replacement is then required which might interrupt surgery. If a patient suffers from tracheostenosis or tracheal

Table 3 Changes of clinical parameters during surgery

Variables	Before anesthesia		During thoracic phase		After thoracic phase	
	SLET group (n=461)	DLET group (n=705) P value	SLET group (n=461)	DLET group (n=705) P value	SLET group (n=461)	DLET group (n=705) P value
HR (bpm)	77.19±11.20	78.24±12.71 0.15	65.21±6.74	65.03±7.26 0.67	61.48±6.82	60.79±6.50 0.08
SBP (mmHg)	118.09±19.37	116.25±18.21 0.10	100.21±12.18	104.04±9.40 <0.01	102.66±13.66	101.87±13.58 0.33
DBP (mmHg)	81.38±15.54	83.79±18.95 0.02	61.97±13.38	65.98±10.75 <0.01	64.62±14.34	63.91±14.98 0.42
SpO ₂ (%)	98.76±0.94	98.82±0.90 0.27	99.71±0.73	98.93±1.16 <0.01	99.81±0.79	99.63±0.85 <0.01
pH	7.40±0.04	7.40±0.04 >0.99	7.30±0.03	7.39±0.05 <0.01	7.38±0.03	7.39±0.05 <0.01
PO ₂ (mmHg)	87.20±8.54	86.77±8.13 0.39	216.89±45.21	170.57±23.16 <0.01	231.67±46.25	230.44±47.80 0.66
PCO ₂ (mmHg)	39.92±3.23	40.15±3.24 0.24	51.17±6.50	40.26±3.13 <0.01	40.20±3.11	39.99±3.19 0.27
PetCO ₂ (mmHg)	-	-	46.07±4.89	35.22±3.17 <0.01	35.00±3.22	35.12±3.15 0.53

SLET, single lumen endotracheal tube; DLET, double lumen endotracheal tube; HR, heart rate; SBP, systolic blood pressure; DBP, diastolic blood pressure.

malformation, application of DLET seems very hard, and sometimes even impossible. The application of SLET, with easy access and management, does not have these defects. SLET with artificial pneumothorax has been adopted for more than 400 cases of MIE in our center. And its exposure of surgical area is as good as DLET (*Figure 1*).

Another defect of one lung ventilation with DLET is the requirement of good pulmonary function (18). Patients with poor pulmonary function might not tolerate one lung ventilation during surgery. Arterial oxygen tension decrease was sometimes observed during surgery with one lung ventilation. Although hypoxic pulmonary vasoconstriction of the nonventilated lung might decrease the hypoxia effect, it's still not as good as two lung ventilation and occasionally develops to severe hypoxemia. This complication is commonly observed in elderly patients with poor pulmonary function (19). Some respiratory complications of MIE are related to one lung ventilation and intraoperative hypoxemia. A few patients would even develop postoperative pulmonary atelectasis and infection after one lung ventilation with DLET. We compared the intraoperative PO₂ level between patients in SLET group and DLET group, and found the incidence of hypoxemia was greatly decreased in SLET group. In the 461 patients in SLET group, none was observed with intraoperative hypoxemia. Statistical analysis also proved significantly higher PO₂ level during thoracic phase in SLET group (*Table 2*).

Safety is the main concern of surgical anesthesia. Artificial pneumothorax with CO₂ insufflation helps exposure the surgical area needed, but also might cause severe complications as reported (7,12), including cardiovascular collapse, ventricular arrhythmias, and contralateral pneumothorax. However, reported cases of severe complications are rare and most of these studies have some limitations, like small amount of enrolled patient, inappropriate pressure of insufflated CO₂, unstandardized protocols, and so on. Further evaluation is necessary. In our study, we retrospectively studied 461 patients underwent artificial pneumothorax with CO₂ as SLET group retrospectively. Eight mmHg of CO₂ insufflation pressure was adopted during thoracic phase of MIE. No severe respiratory and circulatory complications were observed. We compared the circulatory and respiratory data between SLET group and DLET group. In SLET group, slight change of blood pressure and HR could be found, which was induced by 8 mmHg pressure of CO₂ insufflation. Three cases of intraoperative hypotension were observed, but immediately reversed by removal of CO₂ insufflation.

The pH value was greatly decreased, while PO_2 , PCO_2 , and $PetCO_2$ were greatly increased after CO_2 insufflation (Table 3). CO_2 absorption could induce respiratory acidosis and increase the PCO_2 and $PetCO_2$. But these changes did not last long and returned to normal level after removal of CO_2 insufflation. Thus, our research indicates that artificial pneumothorax with CO_2 insufflation is safe under limited pressure of insufflated CO_2 and standardized protocol.

Another possible defect of SLET is the requirement of consistent CO_2 insufflation and stable pressure. When the surgeon decided to change to open surgery or Ivor Lewis esophagectomy during McKeown MIE, it would be hard to maintain good surgical exposure without CO_2 pressure. Under this condition, SLET needs to be changed to DLET during surgery.

In order to evaluate the clinical feasibility of two lung ventilation in MIE, we compared the surgical variables and complications between SLET group and DLET group. With similar background of patient characteristics, SLET group seemed to have shorter total operative time, shorter thoracic phase time, shorter postoperative hospital stay, more harvested left RLN lymph nodes, and more total RLN lymph nodes than DLET group. Our data included MIE cases from 2006 to 2016, but SLET was only applied since 2013. Thus, the better outcome of surgical variables of SLET group might be related to more experienced thoracic surgeon in recent years.

In conclusion, our data presented that two lung ventilation with artificial pneumothorax is a safe and feasible choice for MIE. It has the advantages of simple management, good exposure, few complications, and so on. Circulatory inhibition might be a focus of SLET disadvantages, while our study showed no case of severe circulatory complications under limited CO_2 pressure and standardized protocol.

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Footnote

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

Ethical Statement: This study has been approved by the

Ethics Committee of Zhongshan Hospital (No. B2017-153), and all patients signed informed consent.

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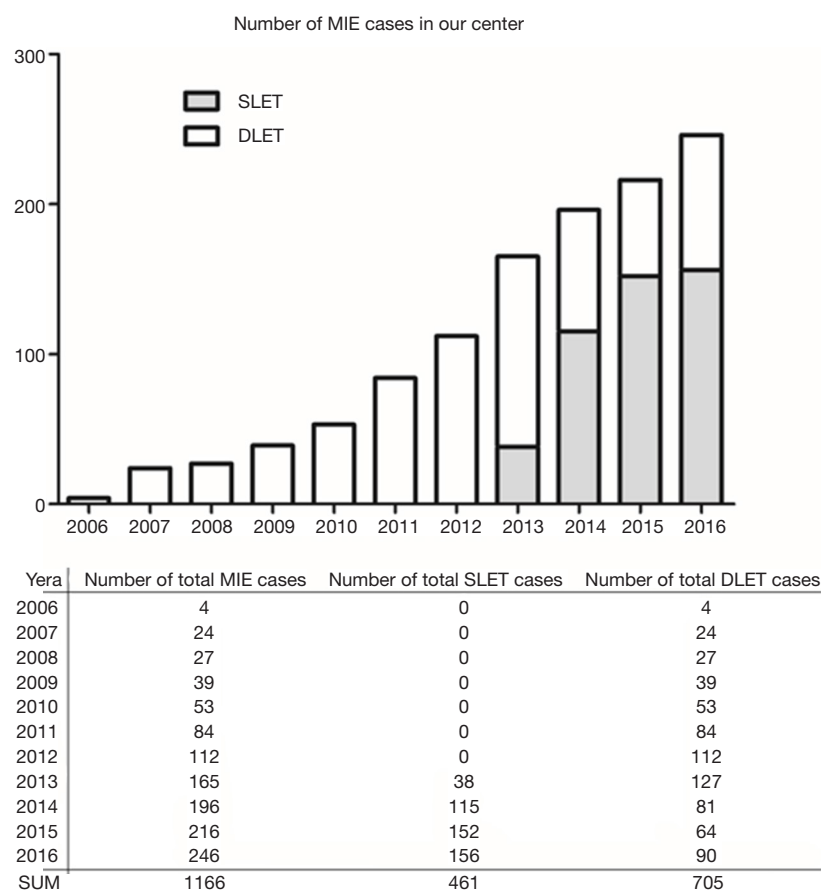


Figure S1 Number of MIE cases in our center from 2006–2016. MIE, minimally invasive esophagectomy; SLET, single lumen endotracheal tube; DLET, double lumen endotracheal tube.