

A Prospective Randomized Trial of Enteral Nutrition After Thoracoscopic Esophagectomy for Esophageal Cancer

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ABSTRACT

Background. Several studies have reported that postoperative enteral nutrition (EN) reduced complications and decreased weight loss and hospital stay periods; however, the majority of patients analyzed in these studies underwent open thoracic surgery. No studies have been conducted regarding EN in patients after thoracoscopic esophagectomy as a less invasive surgery. The aim of this study was to investigate the efficacy of EN after thoracoscopic esophagectomy.

Methods. Fifty patients who underwent thoracoscopic esophagectomy for esophageal cancer were divided into two groups: parenteral nutrition (PN; $n = 25$) and EN ($n = 25$). The rate of weight loss at postoperative day (POD) 14, levels of prealbumin at POD 10, postoperative complications until POD 14, and other perioperative data were collected for each group.

Results. This study analyzed data for 47 patients. The rate of weight loss at POD 14 was significantly lower in the EN group ($3.0 \pm 3.2\%$) than in the PN group ($4.0 \pm 3.6\%$; $p = 0.020$). Prealbumin levels were 21.0 ± 7.5 mg/dL in the PN group and 18.4 ± 5.8 mg/dL in the EN group at POD 10, with no significant differences between the groups. However, the incidence of postoperative pneumonia was higher in the PN group (30.4%) than in the EN group (12.5%).

Conclusions. EN could suppress weight loss and reduce the incidence of pneumonia after thoracoscopic esophagectomy.

Esophageal cancer is a highly aggressive malignancy that metastasizes to the lymph nodes and is associated with a poor prognosis. The 5-year overall survival rate is 40.0% and the 30-day mortality rate is 1.7%.¹ Surgical resection is the most effective treatment for localized esophageal cancer; however, esophagectomy is extremely invasive and is associated with high morbidity and mortality rates.

Thoracoscopic esophagectomy for esophageal cancer was first reported by Cuschieri et al.² in 1992 and has been increasingly adopted as a minimally invasive esophagectomy. A current review of thoracoscopic esophagectomy showed that single-institution studies and several meta-analyses have demonstrated acceptable short-term outcomes compared with open esophagectomy.³ The first multicenter trial comparing thoracoscopic esophagectomy and open esophagectomy [TIME trial (Traditional Invasive vs. Minimally Invasive Esophagectomy)] reported that the incidence of postoperative pulmonary infection was markedly lower in the thoracoscopic group. Additional benefits of thoracoscopic esophagectomy in this trial were less operative blood loss, better postoperative quality of life for patients, and shorter hospital stay.⁴

Nutrition is one of the most important factors to consider after esophagectomy in order to reduce surgical mortality.^{5,6} The European Society for Parenteral and Enteral Nutrition guidelines recommend early tube feeding after major gastrointestinal surgery for cancer.⁷ Several studies have shown that enteral nutrition (EN) is more effective than parenteral nutrition (PN) in reducing postoperative complications in postesophagectomy patients,^{8–10} however, these studies mainly analyzed patients who underwent open esophagectomy. No studies have been conducted showing the efficacy of EN in patients after thoracoscopic esophagectomy.

We hypothesized that EN was also recommended after thoracoscopic esophagectomy in esophageal cancer

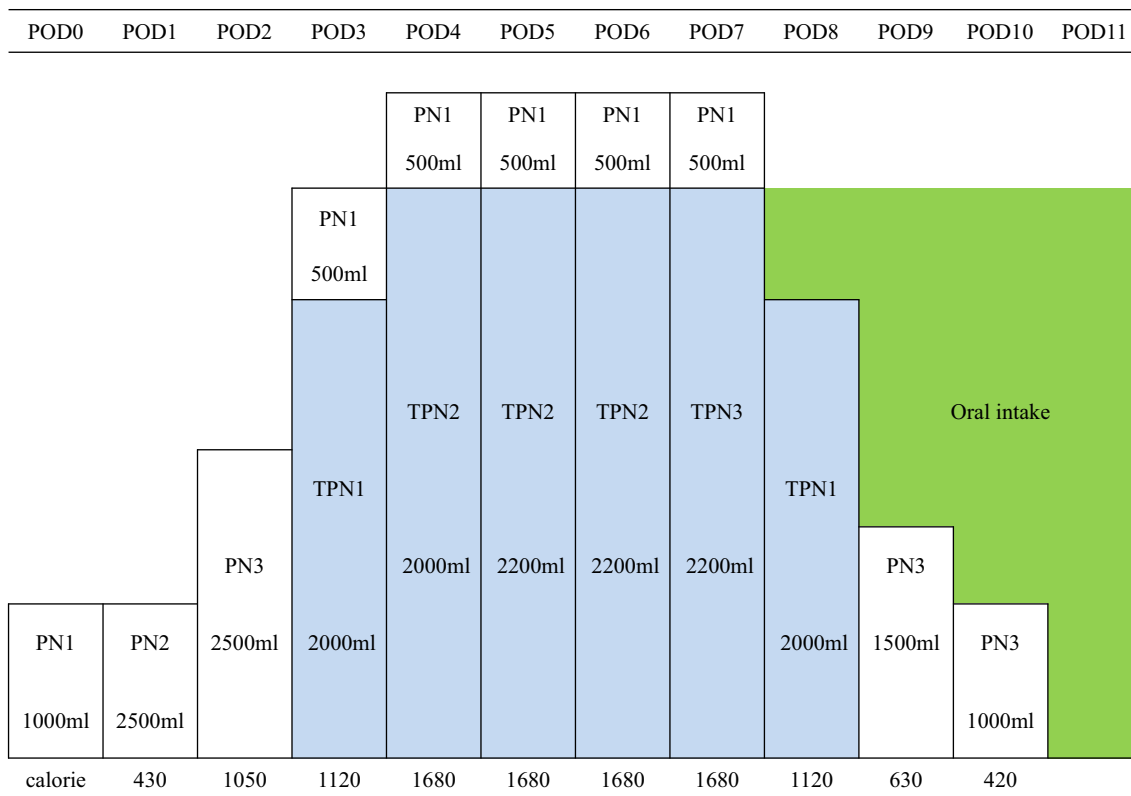


FIG. 1 Protocol for parenteral nutrition. PN1: sodium 130 mEq/L, potassium 4 mEq/L, calcium 3 mEq/L, chlorine 109 mEq/L, lactate 28 mEq/L, glucose 0 g/L, calories 0 kcal/L; PN2: sodium 35 mEq/L, potassium 20 mEq/L, chlorine 35 mEq/L, lactate 20 mEq/L, glucose 43 g/L, calories 172/L; PN3: sodium 35 mEq/L, potassium 20 mEq/L, magnesium 5 mEq/L, calcium 5 mEq/L, chlorine 35 mEq/L, sulfate 5 mEq/L, lactate 20 mEq/L, acetate 16 mEq/L, glucose 75 g/L, free amino acid 30 g/L, thiamine 1.5 mg, calories 420/L; TPN1: sodium

50 mEq/L, potassium 22 mEq/L, magnesium 4 mEq/L, calcium 4 mEq/L, chlorine 50 mEq/L, sulfate 4 mEq/L, lactate 12 mEq/L, acetate 41 mEq/L, glucose 120 g/L, free amino acid 20 g/L, calories 560/L; TPN2: sodium 50 mEq/L, potassium 27 mEq/L, magnesium 5 mEq/L, calcium 5 mEq/L, chlorine 50 mEq/L, sulfate 5 mEq/L, lactate 15 mEq/L, acetate 50 mEq/L, glucose 175 g/L, free amino acid 30 g/L, calories 820/L. *POD* postoperative day

patients. In this study, we conducted a randomized controlled trial comparing PN and EN after thoroscopic esophagectomy.

MATERIALS AND METHODS

After thoroscopic esophagectomy for esophageal cancer, patients were randomly divided into the PN and EN groups and treated as per each study protocol. The medical data of the patients were prospectively collected. The primary endpoint was the rate of weight loss at postoperative day (POD) 14 from preoperative weight, while secondary endpoints were serum prealbumin levels at POD 10 and postoperative complications until POD 14.

Patient selection criteria were (i) patients undergoing planned thoroscopic esophagectomy with two- or three-field lymphadenectomy at Keio University Hospital, Tokyo, Japan; (ii) patient consent to participate in this study; (iii) age 20–75 years; (iv) body mass index (BMI) 18–25 kg/m²; and (v) an American Society of Anesthesiologists physical

status of I or II. Exclusion criteria were severe ischemic heart disease, cerebrovascular disease, chronic renal failure (creatinine clearance rate <30 mL/min), severe diabetes, liver dysfunction, severe obesity (BMI >25 kg/m²), severe emaciation (BMI <18 kg/m²), amino acid metabolism disorder, pregnancy, and preoperative chemoradiation therapy.

Patients in the PN group received total PN from a central venous catheter, as per the study protocol (Fig. 1), and patients in the EN group underwent jejunostomy during esophagectomy, and commenced 24-h continuous jejunostomy feeding of an elemental diet immediately after surgery. According to the study protocol (Fig. 2), the initial dose of the enteral diet was 10 mL/h, gradually increasing to 70 mL/h. Two study protocols were designed to comprise almost identical total calories and total moisture content.

Although patients essentially received nutrition as per either protocol, in both groups, the quantity of infusion solutions or enteral feeding solutions was changed depending on the patient’s condition. For example, an additional transfusion to improve dehydration or a

reduction in enteral solution due to diarrhea was included. Total intake quantity and total calories were retrospectively calculated for each patient.

All patients consumed an enteral diet of 400 mL/day orally for 5 days prior to surgery. A peripherally inserted central catheter was inserted from the arm before surgery, and hydrocortisone (200 mg/day) was administered for 5 days—2 days before and 3 days after surgery.

During surgery, the thoracic procedure was performed by video-assisted thoracoscopic surgery, and the abdominal procedure was performed by open laparotomy or hand-assisted laparoscopic surgery (HALS), as previously described.¹¹ Fields of lymph node dissection and thoracic duct resection were decided according to the tumor progression and preoperative risk of each patient. A single surgical team including three operators performed all the thoracoscopic esophagectomies, in which two of the three surgeons always participated as assistants.

After surgery, patients were treated with an artificial ventilator in the intensive care unit (ICU), until POD 1. Patients routinely underwent computed tomography (CT) scans to detect postoperative complications on POD 6, and contrast imaging to check swallowing functions on POD 7. As a result of the CT scan and swallowing examination, patients without severe complications started oral intake on POD 8. Thereafter, the quantity of infusion solutions or enteral feeding solutions was decreased depending on oral intake. Patients who recovered well were discharged after POD 14. Blood sugar levels were measured four times a day in all patients, and insulin was injected or mixed into infusion solutions, depending on these levels.

Collected postoperative data included weight at POD 14, prealbumin level at POD 10, postoperative complications until POD 14, oral intake commencement day, ICU stay and postoperative hospital-stay durations, total intake (infusion solution and enteral diet) and discharge (urine and drain discharge) quantities until POD 14, and postoperative blood test results [total protein, albumin, total bilirubin, aspartate transaminase, alanine aminotransferase, blood sugar, C-reactive protein (CRP) levels, white blood cell (WBC) and platelet counts, and prothrombin time]. Postoperative complications were diagnosed using X-ray, computed tomography, and clinical features, and the severity of complications was classified using the Clavien–Dindo classification grade of surgical complications.¹²

Statistical analyses between the groups were performed using the Mann–Whitney *U* test and Chi square test using IBM SPSS statistical software version 22 (IBM Corporation, Armonk, NY, USA). A probability (*p*) value of <0.05 was considered statistically significant.

Sample size was determined on the basis of our previous postoperative data of 52 patients administered EN or PN

after esophagectomy during 2006–2009. The mean rate of weight loss from preoperative weight at POD 14 was 6.0 % in the EN group and 7.5 % in the PN group. Calculated sample size for which 80 % statistical power could be expected was 34 patients (17 patients/group), on the assumption that the standard deviation was 5 and mean weight loss rate improved by 5 % using EN. The sample size of this study was determined as 50 patients (25 patients/group) to allow for accidental errors.

The study protocol was approved by the Ethics Committee of Keio University School of Medicine (UMIN000014884), and written informed consent was obtained from all patients in the study.

RESULTS

Patient Characteristics

Fifty patients were enrolled in this study from March 2012 to July 2014, and the data of 47 patients were analyzed. Three patients were excluded from analysis because the surgery was converted to open surgery in one patient, one patient in the PN group received enteral feeding because of severe anastomotic leakage, and one patient in the EN group received only PN because of additional surgery (colectomy) for another disease.

Patient characteristics are shown in Table 1. The 47 patients analyzed comprised 37 (78.8 %) men and 10 (21.2 %) women, with a mean age of 62.2 years (range 46–74). There were no significant differences in age, sex, height, weight, medical history, preoperative blood test results, neoadjuvant chemotherapy, or pathological TNM classification (UICC TNM Classification of Malignant Tumors, 7th edition).

Surgical Procedures

Intraoperative data are summarized in Table 2. The abdominal procedure was performed with HALS in 46 (97.9 %) patients. Forty-three (95.7 %) patients underwent three-field lymph node dissection and 45 (95.7 %) patients underwent thoracic duct resection. Mean surgical duration was 556 ± 52 min and mean blood loss was 208 ± 139 mL. No significant differences in the surgical procedure between the two groups were observed.

Primary and Secondary Endpoint

The results of primary and secondary endpoints are shown in Table 3. The mean rate of weight loss for all patients at POD 14 was 4.0 ± 3.6 : 5.1 ± 3.7 % in the PN group and 3.0 ± 3.2 % in the EN group. Weight loss after

TABLE 1 Characteristics of patients

	All patients (n = 47)		Parenteral nutrition (n = 23)		Enteral nutrition (n = 24)		p value
Age (years)	62.2 ± 8.12		60.7 ± 8.97		63.6 ± 7.13		0.281
Sex (M/F)	37/10		18/5		19/5		0.940
Height (m)	1.66 ± 0.10		1.66 ± 0.12		1.66 ± 0.06		0.790
Weight (kg)	59.8 ± 11.9		57.2 ± 12.1		62.3 ± 11.3		0.225
Medical history [n (%)]							
Hypertension	13 (27.7)		4 (17.4)		9 (37.5)		0.127
Diabetes mellitus	5 (10.6)		1 (4.3)		4 (16.7)		0.176
Malignant tumor	3 (6.4)		1 (4.3)		2 (8.3)		0.580
Hyperlipidemia	3 (6.4)		0 (0.0)		3 (12.5)		0.083
Gastric ulcer	3 (6.4)		3 (13.0)		0 (0.0)		0.070
Hepatitis	2 (4.3)		1 (4.3)		1 (4.2)		0.976
COPD	1 (2.1)		1 (4.3)		0 (0.0)		0.307
Others	5 (10.6)		2 (8.7)		3 (12.5)		0.676
Preoperative blood test							
Total protein (g/dL)	6.4 ± 0.4		6.4 ± 0.4		6.4 ± 0.5		0.923
Albumin (g/dL)	4.0 ± 0.4		4.1 ± 0.2		4.0 ± 0.4		0.708
Total bilirubin (mg/dL)	0.7 ± 0.2		0.7 ± 0.2		0.7 ± 0.3		0.400
AST (U/L)	23 ± 10		21 ± 8.1		24 ± 11		0.281
ALT (U/L)	20 ± 17		17 ± 13		23 ± 19		0.141
HbA1c (%)	5.8 ± 0.6		5.8 ± 0.5		5.8 ± 0.8		0.543
CRP (mg/dL)	0.11 ± 0.21		0.13 ± 0.26		0.09 ± 0.14		0.889
WBC (×10 ³ /μL)	6.4 ± 2.2		6.6 ± 2.4		6.3 ± 2.1		0.709
Plt (×10 ⁴ /μL)	22.9 ± 6.2		23.5 ± 5.7		22.3 ± 6.7		0.273
Prothrombin time (%)	96.4 ± 7.7		97.7 ± 6.1		95.2 ± 9.0		0.211
Preoperative therapy [n (%)]							
Chemotherapy	25	(53.2)	13	(56.5)	12	(50.0)	0.658
TNM classification (UICC 7th edition) [n (%)]							
Primary tumor							0.643
pT1	34	(72.3)	16	(69.6)	18	(75.0)	
pT2	4	(8.5)	3	(13.0)	1	(4.2)	
pT3	9	(19.1)	4	(17.4)	5	(20.8)	
Resional lymph node							0.493
pN0	25	(53.2)	13	(56.5)	12	(50.0)	
pN1	16	(34.0)	8	(34.8)	8	(33.3)	
pN2	3	(6.4)	2	(8.7)	1	(4.2)	
pN3	3	(6.4)	0	(0.0)	3	(12.5)	
Distant metastasis							0.418
pM0	41	(87.2)	21	(91.3)	20	(83.3)	
pM1 (lymph node metastasis)	6	(12.8)	2	(8.7)	4	(16.7)	
Stage							0.469
I	21	(44.7)	11	(47.8)	10	(41.7)	
II	13	(27.7)	7	(30.4)	6	(25.0)	
III	7	(14.9)	3	(13.0)	4	(16.7)	
IV	6	(12.8)	2	(8.7)	4	(16.7)	

COPD chronic obstructive pulmonary disease, AST aspartate transaminase, ALT alanine aminotransferase, CRP C-reactive protein, WBC white blood cell, M male, F female, UICC Union for International Cancer Control, ± indicates mean ± SD

TABLE 2 Intraoperative data

	All patients (<i>n</i> = 25)	Parenteral nutrition (<i>n</i> = 23)	Enteral nutrition (<i>n</i> = 24)	<i>p</i> -Value
Surgical procedure [<i>n</i> (%)]				
Hand-assisted laparoscopic surgery	46 (97.9)	23 (100)	23 (95.8)	0.328
Lymph node dissection				
Two-fields	4 (8.5)	1 (4.3)	3 (12.5)	0.322
Three-fields	43 (91.5)	22 (95.7)	21 (87.5)	
Number of dissected lymph nodes (mean ± SD)	72 ± 23	67 ± 26	77 ± 19	0.053
Thoracic duct resection [<i>n</i> (%)]	45 (95.7)	22 (95.6)	23 (95.8)	0.976
Operation time (min; mean ± SD)	556 ± 52	548 ± 53	564.2 ± 51.2	0.395
Blood loss (mL; mean ± SD)	208 ± 139	211 ± 122	207 ± 157	0.616
Blood transfusion (mL; mean ± SD)	88 ± 222	36 ± 128	137 ± 279	0.263
Albumin solution (mL; mean ± SD)	1044 ± 299	989 ± 324	1097 ± 269	0.155

SD standard deviation

surgery was significantly prevented in the EN group ($p = 0.020$).

Mean prealbumin level was 19.7 ± 6.8 mg/dL at POD 10: 21.0 ± 7.5 mg/dL in the PN group and 18.4 ± 5.8 mg/dL in the EN group. No significant differences were observed between the groups ($p = 0.257$).

Postoperative complications occurred in 34 (72.3 %) of 47 patients. Severe complications (Clavien–Dindo grade 3 or higher) occurred in eight (17.0 %) patients: four with recurrent nerve paralysis, three with pneumonia, one with cerebral hemorrhage, and one with thoracic emphysema. No significant differences were observed between groups for all complications ($p = 0.680$), each complication, or severe complications ($p = 0.137$). However, the incidence of pneumonia was higher in the PN group (30.4 %, seven patients) than in the EN group (12.5 %, three patients). One patient in each group had a catheter-associated infection. In the EN group, no severe complication relating to enteral feeding, such as peritonitis, ileus, or severe diarrhea, was observed. In both groups, the postoperative mortality rate was 0 %.

Postoperative Data

Postoperative data are shown in Table 3. Mean ICU stay was 2.7 days in both groups ($p = 0.327$). The mean PODs of oral intake commencement were 13.9 ± 7.7 in the PN group and 15.1 ± 9.5 in the EN group ($p = 0.664$). Postoperative hospital stay was longer in the EN group (30.5 ± 9.0 days) than in the PN group (27.8 ± 14.3 days). No differences were observed in durations of ICU stay, oral intake, and hospital stay ($p = 0.327$, $p = 0.664$, and $p = 0.058$, respectively).

Postoperative blood test results were almost identical in the two groups. No significant differences ($p = 0.912$) were observed in the mean values of blood sugar levels during the

10-day period after surgery— 155.2 ± 21.3 mg/dL in the PN group and 159.7 ± 28.6 mg/dL in the EN group.

During the 7-day period after surgery, no differences in the mean values of intake (infusion solution and enteral diet) and calories were observed. However, the mean amount of discharge (urine and drain discharge) was significantly larger in the PN group (2.9 L/day) than in the EN group (2.5 L/day) ($p = 0.011$).

Five patients in the EN group required enteral feeding to be continued after discharge from hospital in order to support their nutrition. Two patients in the PN group, complicated by cerebral hemorrhage or severe recurrent laryngeal nerve paralysis, required enteral feeding, using a nasojunal tube, on POD 17 and POD 28.

DISCUSSION

To the best of our knowledge, this is the first prospective randomized study of early EN after thoracoscopic esophagectomy. The results of this study demonstrated that postoperative EN suppressed weight loss at POD 14. The incidence of pneumonia tended to be lower in the PN group than in the EN group, and no significant differences were observed in serum levels of prealbumin, or incidence of complications, between the PN and EN groups.

Several studies have advocated that after esophagectomy, postoperative EN reduces the incidence of complications,^{8–10} duration of ICU stay,¹³ duration of hospital stay,^{9,13} and postoperative weight loss.¹⁴ Early recovery of the total WBC count and decrease in the serum levels of total bilirubin and CRP in patients with early EN have also been reported.¹⁵

In this study, weight loss after surgery was significantly prevented in the EN group. Although no differences in administered calories or moisture content were observed, the amount of discharge until POD 7 was significantly

TABLE 3 Postoperative data

	All patients (<i>n</i> = 47)	Parenteral nutrition (<i>n</i> = 23)	Enteral nutrition (<i>n</i> = 24)	<i>p</i> -Value
Primary endpoint				
Rate of weight loss at 14 POD (%; mean ± SD)	4.00 ± 3.56	5.05 ± 3.65	2.94 ± 3.19	0.020
Secondary endpoint				
Prealbumin at 10 POD (mg/dL; mean ± SD)	19.7 ± 6.77	21.0 ± 7.48	18.4 ± 5.82	0.257
Complications [<i>n</i> (%)]				
Recurrent laryngeal nerve paralysis	13 (27.7)	6 (26.1)	7 (29.2)	0.815
Pneumonia	10 (21.3)	7 (30.4)	3 (12.5)	0.137
Anastomotic leakage	8 (17.0)	4 (17.4)	5 (20.8)	0.767
Thrombosis	7 (14.9)	4 (17.4)	3 (12.5)	0.641
Atelectasis	4 (8.5)	2 (8.7)	2 (8.3)	0.965
Lymphorrhoea	4 (8.5)	2 (8.7)	2 (8.3)	0.965
Others	9 (19.1)	4 (17.4)	5 (20.8)	0.767
Any complications	34 (72.3)	16 (69.6)	18 (75.0)	0.680
Severe complications	8 (17.0)	5 (21.7)	3 (12.5)	0.461
Postoperative data (mean ± SD)				
Nutrition and balance (until POD 7)				
Calories of intake (kcal/day)	1103.0 ± 273.2	1104.3 ± 202.7	1101.6 ± 159.0	0.483
Intake (mL/day)	3011.1 ± 322.0	2985.2 ± 345.8	3036.0 ± 302.7	0.407
Discharge (mL/day)	2708.9 ± 502.4	2891.1 ± 481.45	2534.3 ± 467.0	0.011
Postoperative course				
Start of oral intake (POD)	14.5 ± 8.7	13.9 ± 7.7	15.1 ± 9.5	0.664
ICU stay (days)	2.7 ± 2.0	2.7 ± 2.3	2.7 ± 1.6	0.327
Postoperative hospital stay (days)	27.7 ± 11.8	27.1 ± 14.7	28.3 ± 8.4	0.147
Postoperative blood test				
Total protein (at POD14; g/dL)	5.9 ± 0.4	6.0 ± 0.4	5.9 ± 0.6	0.811
Albumin (at POD14; g/dL)	3.1 ± 0.4	3.2 ± 0.3	3.1 ± 0.5	0.57
Total bilirubin (mg/dL)	1.7 ± 1.1	2.0 ± 1.4	1.3 ± 0.7	0.079
AST (maximal values; IU/L)	66.2 ± 43.5	58.0 ± 21.1	74.1 ± 56.7	0.882
ALT (maximal values; IU/L)	83.8 ± 74.8	74.3 ± 45.2	92.8 ± 95.3	0.717
Blood sugar (mean of 10 days; mg/dL)	157.5 ± 25.3	155.2 ± 21.5	159.7 ± 28.6	0.912
CRP (maximal values; mg/mL)	13.4 ± 4.0	13.3 ± 4.0	13.5 ± 4.1	0.406
WBC (maximal values; ×10 ³ /μL)	13.4 ± 4.0	13.6 ± 4.3	13.2 ± 3.8	0.595
Platelet (minimal values; ×10 ⁴ /μL)	15.3 ± 4.2	15.4 ± 4.5	15.3 ± 4.1	0.941
Prothrombin time (minimal values; %)	68.0 ± 11.5	71.0 ± 8.5	67.0 ± 10.5	0.338

AST aspartate transaminase, ALT alanine aminotransferase, CRP C-reactive protein, WBC white blood cell, SD standard deviation, POD postoperative day, ICU intensive care unit

Thrombosis: pulmonary embolism, deep vein thrombosis, or jugular vein thrombosis

Severe complications: complication diagnosed as Clavien–Dindo grade 3 or higher

smaller in the EN group. In the PN group, more moisture may have been transferred to the third space or urine than in the EN group. EN may also superiorly help in absorbing nutrients and suppressing weight loss compared with PN. However, no differences in prealbumin levels at POD 10 were observed between the two groups. Prealbumin is known as the earliest laboratory indicator of nutritional status.¹⁶ Because prealbumin is also a marker of tissue permeability, it is affected by inflammatory response and

steroid use.¹⁷ Prealbumin cannot exactly reflect nutritional status after invasive surgery.

EN and postoperative complications, particularly infectious complications, after gastrointestinal surgery are related.^{6,8} Few mechanisms whereby EN reduced postoperative complications have been reported. EN prevents atrophy of gastrointestinal mucosa and inhibits bacterial translocation from the gut to the blood stream.¹⁸ It reduces sequestration of the fluid in the third space and improves pulmonary

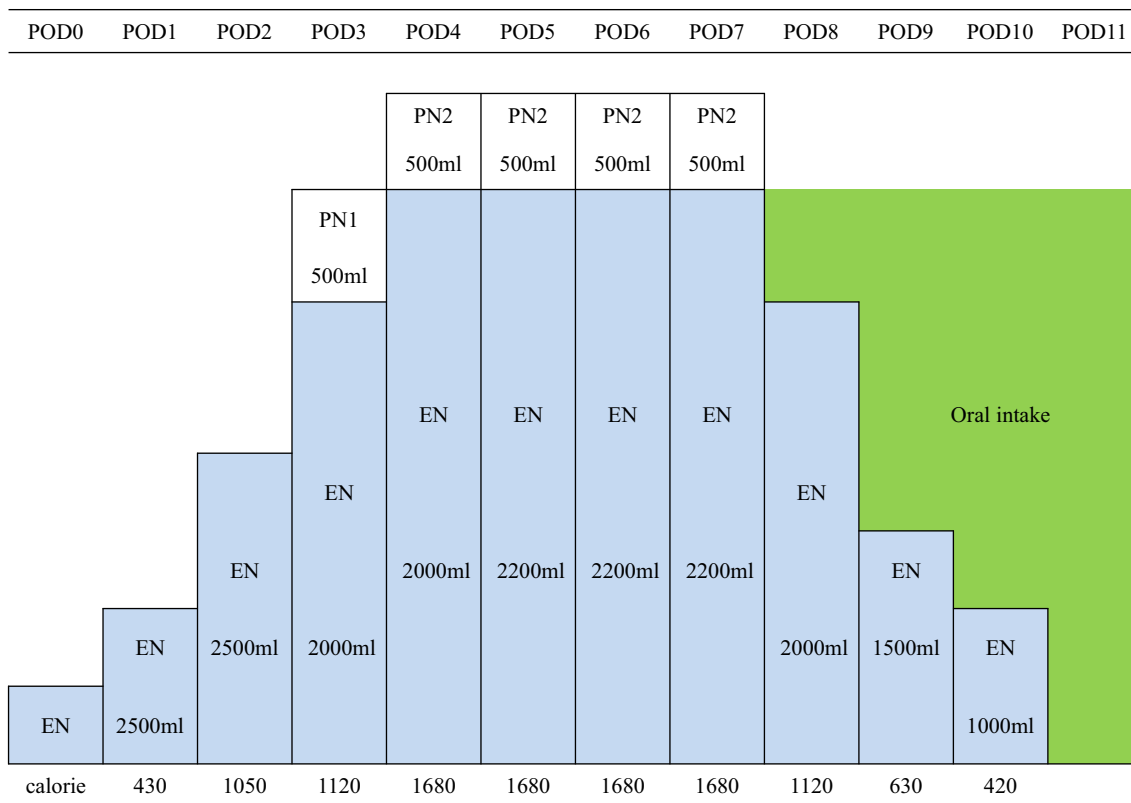


FIG. 2 Protocol for enteral nutrition. EN1: 1 kcal/1 mL, osmotic pressure 760 mOsm/L (free amino acid 44 g/L, carbohydrate 211 g/L, lipid 1.7 g/L). *POD* postoperative day

function.¹⁹ Moreover, EN normalizes the immune system.²⁰ In this study, the incidence of pneumonia was less in the EN group. EN may also reduce infectious complications in patients after thoracoscopic esophagectomy. To clarify the effect of EN for postoperative complications, a greater number of patients need to be analyzed.

Unlike the previous study, no differences were observed in total bilirubin and CRP levels, and duration of ICU and postoperative hospital stay, in this study. For patients who underwent the thoracoscopic procedure and steroid therapy to reduce excessive inflammatory response and related complications, enteral feeding did not greatly contribute to their recovery.

Hyperglycemia is one of the most important complications of PN.²¹ In our study, for the five patients with mild to moderate diabetes mellitus, the mean blood sugar levels until POD 14 were almost equivalent among the two groups. These results suggested that blood sugar could be controlled adequately by sliding-scale insulin therapy in both groups.

In seven patients, enteral feeding was needed after discharge. Four patients had dysphagia with recurrent laryngeal nerve paralysis, two experienced loss of appetite, and one had cerebral hemorrhage after esophagectomy. EN

was particularly beneficial for patients with insufficient oral intake even after discharge.

CONCLUSIONS

Postoperative EN was recommended after thoracoscopic esophagectomy to suppress both postoperative weight loss and the incidence of pneumonia. EN was also beneficial for patients with insufficient oral intake postsurgery.

DISCLOSURE This study was not supported by any funding sources.

REFERENCES

1. Ando N, Ozawa S, Kitagawa Y, et al. Improvement in the results of surgical treatment of advanced squamous esophageal carcinoma during 15 consecutive years. *Ann Surg.* 2000;232:225–32.
2. Cuschieri A, Shimi S, Banting S. Endoscopic oesophagectomy through a right thoracoscopic approach. *J R Coll Surg Edinb.* 1992;37:7–11.
3. Takeuchi H, Kawakubo H, Kitagawa Y. Current status of minimally invasive esophagectomy for patients with esophageal cancer. *Gen Thorac Cardiovasc Surg.* 2013;61:513–21.
4. Biere SS, van Berge Henegouwen MI, Maas KW, et al. Minimally invasive versus open oesophagectomy for patients with

- oesophageal cancer: a multicentre, open-label, randomised controlled trial. *Lancet*. 2012;379:1887–92.
5. Lewis SJ, Andersen HK, Thomas S. Early enteral nutrition within 24 h of intestinal surgery versus later commencement of feeding: a systematic review and meta-analysis. *J Gastrointest Surg*. 2009;13:569–75.
 6. Zhu XH, Wu YF, Qiu YD, et al. Effect of early enteral combined with parenteral nutrition in patients undergoing pancreaticoduodenectomy. *World J Gastroenterol*. 2013;19:5889–96.
 7. Weimann A, Braga M, Harsanyi L, et al. ESPEN guidelines on enteral nutrition: surgery including organ transplantation. *Clin Nutr*. 2006;25:224–44.
 8. Baigrie RJ, Devitt PG, Watkin DS. Enteral versus parenteral nutrition after oesophagogastric surgery: a prospective randomized comparison. *Aust N Z J Surg*. 1996;66:668–70.
 9. Fujita T, Daiko H, Nishimura M. Early enteral nutrition reduces the rate of life-threatening complications after thoracic esophagectomy in patients with esophageal cancer. *Eur Surg Res*. 2012;48:79–84.
 10. Couper G. Jejunostomy after esophagectomy: a review of evidence and current practice. *Proc Nutr Soc*. 2011;70:316–20.
 11. Kaburagi T, Takeuchi H, Kawakubo H, et al. Clinical utility of a novel hybrid position combining the left lateral decubitus and prone positions during thoracoscopic esophagectomy. *World J Surg*. 2014;38:410–8.
 12. Dindo D, Demartines N, Clavien PA. Classification of surgical complications: a new proposal with evaluation in a cohort of 6336 patients and results of a survey. *Ann Surg*. 2004;240:205–13.
 13. Gabor S, Renner H, Matzi V, et al. Early enteral feeding compared with parenteral nutrition after oesophageal or oesophagogastric resection and reconstruction. *Br J Nutr*. 2005;93:509–13.
 14. Watters JM, Kirkpatrick SM, Norris SB, et al. Immediate postoperative enteral feeding results in impaired respiratory mechanics and decreased mobility. *Ann Surg*. 1997;226:369–77; (discussion 377–80).
 15. Aiko S, Yoshizumi Y, Sugiura Y, et al. Beneficial effects of immediate enteral nutrition after esophageal cancer surgery. *Surg Today*. 2001;31:971–8.
 16. Beck FK, Rosenthal TC. Prealbumin: a marker for nutritional evaluation. *Am Fam Physician*. 2002;65:1575–8.
 17. Sodergren MH, Jethwa P, Kumar S, et al. Immunonutrition in patients undergoing major upper gastrointestinal surgery: a prospective double-blind randomised controlled study. *Scand J Surg*. 2010;99:153–61.
 18. Feng Y, Ralls MW, Xiao W, et al. Loss of enteral nutrition in a mouse model results in intestinal epithelial barrier dysfunction. *Ann N Y Acad Sci*. 2012;1258:71–7.
 19. Austrums E, Pupelis G, Snippe K. Postoperative enteral stimulation by gut feeding improves outcomes in severe acute pancreatitis. *Nutrition*. 2003;19:487–91.
 20. Okada Y, Klein N, van Saene HK, et al. Small volumes of enteral feedings normalise immune function in infants receiving parenteral nutrition. *J Pediatr Surg*. 1998;33:16–9.
 21. Gosmanov AR, Umpierrez GE. Management of hyperglycemia during enteral and parenteral nutrition therapy. *Curr Diab Rep*. 2013;13:155–62.