

## Does epidural analgesia modify the risk of complications after gastrectomy?

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**Introduction.** The surgical treatment of Gastric Cancer is associated with overall complication rates as high as 50%. The intent of this study was to assess the impact of epidural analgesia (EA) on postoperative complication rates among patients undergoing gastric resections.

**Materials and methods.** Of the 617 gastric cancer patients who between 2002 and 2010 had undergone stomach resection, 246 (39.8%) were administered EA. Groups with and without EA were compared.

**Results.** The general rate of complications was lower in the EA group in the univariable analysis – 38.5% vs. 54.2% (odds ratio [OR]: 0.47, 95% confidence interval [CI]: 0.34–0.66,  $p < 0.001$ ), intra-abdominal abscess (OR 0.28, 95% CI: 0.14–0.59,  $p = 0.001$ ), pneumonia (OR 0.39, 95% CI: 0.24–0.63,  $p < 0.001$ ), temperature  $>38^{\circ}\text{C}$  (OR 0.53, 95% CI: 0.37–0.74,  $p < 0.001$ ) and re-operation (OR 0.53, 95% CI: 0.28–1.00,  $p = 0.049$ ). These relationships were confirmed in a multivariable analysis for the general number of complications (OR 0.53, 95% CI: 0.37–0.75,  $p < 0.001$ ), intra-abdominal abscess (OR 0.36, 95% CI: 0.16–0.77,  $p = 0.009$ ), temperature  $>38^{\circ}\text{C}$  (OR 0.56, 95% CI: 0.39–0.82,  $p = 0.009$ ), pneumonia (OR 0.42, 95% CI: 0.25–0.71,  $p = 0.001$ ).

**Conclusions.** Our findings indicate that postoperative treatment with EA for patients undergoing stomach resection is safe and contributes to a reduction in the number of postoperative complications.

**Key words:** gastric cancer, GEJ cancer, epidural analgesia, postoperative complications, gastrectomy, postoperative pain

### Introduction

For the past 100 years, cases of stomach cancer (gastric cancer – GC) amongst developed countries have been systemically in decline. Possible contributing factors for this decline may be attributed to the increased use of refrigeration for food storage, dietary changes, and decreased incidents of infections with *Helicobacter pylori* [1]. Despite progress, stomach cancer remains the fourth

most frequently diagnosed cancer worldwide. In 2008, there were 980,000 new cases, of which, 83,000 were reported in the European Union and over 5000 in Poland alone [1, 2]. In Poland, the 5-year survival rate post stomach cancer diagnosis is about 18%, in Europe is about 25%, while in Japan about 70% [2–4].

Surgical resection of gastric cancer has produced suboptimal survival rates despite multidisciplinary treatment appro-

### How to cite:

Olesiński T, Urbańska M, Borkowska A, Wieszczy P, Król D, Rucz D, Symonides M. *Does epidural analgesia modify the risk of complications after gastrectomy?* NOWOTWORY J Oncol 2021; 71: 202–211.

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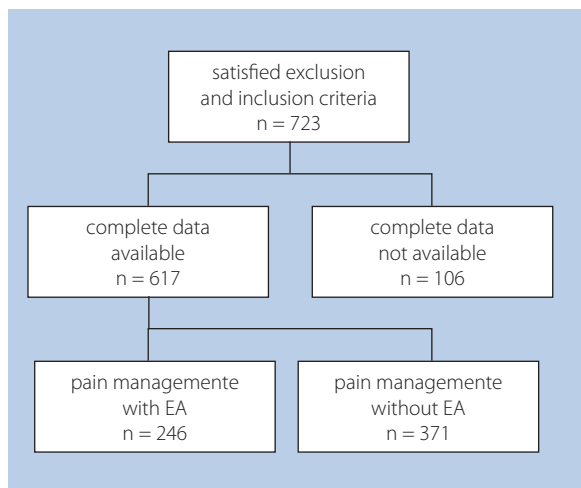
aches and improvements in surgical techniques. The European Society for Medical Oncology (ESMO) guidelines of treatment for patients diagnosed with an advanced GC include peri-operative chemotherapy [5]. However, a total or subtotal gastrectomy with removal of the surrounding lymph nodes, (D2 resection) remains the only curative method of treatment [5–8]. The vast extent of surgical intervention is one of the main contributing factors to the high risk of complications associated with the procedure. The estimated number of complications varies between 17 and 48%. Additionally, gastric resection in conjunction with splenectomy or spleno-pancreatectomy significantly increases the potential for complications [9–14]. The management of quality care in postsurgical settings that include administration of regional analgesia contributes to better treatment outcomes [15, 16]. Due to the limitations in the use of opioids, resulting from the recommendations of the ERAS protocol, the effectiveness of epidural analgesia (EA) is very important. Currently, EA is a standard procedure in our team and for this reason historical data were compared. On the other hand, surgical procedures did not undergo significant modification during the period under review.

## Objective

To assess the impact of EA on postoperative complication rates in patients undergoing subtotal or total gastrectomy for gastric cancer.

## Materials and methods

This study was conducted in a single institution using its administrative database. All patients were treated between 2002 and 2010 at The Maria Skłodowska-Curie Institute, Oncology Center in Warsaw. No neoadjuvant therapy was administered in the analyzed period. Of 723 GC resections performed, 617 cases had complete medical documentation that was adequate for retrospective review (study flow – fig. 1).



**Figure 1.** Flow-chart of the study

**Table I.** Baseline characteristics – demographic data, nutrition status, and comorbidities

Characteristic	Epidural analgesia (EA)		p value
	No n = 371 (%)	Yes n = 246 (%)	
gender:			
• female	119 (58.3)	85 (41.7)	0.522
• male	252 (61)	161 (39)	
age: (median) (IQR)	64 (53–71)	61 (54–72)	0.144
BMI			
• <19	31 (70.5)	13 (29.5)	0.050
• 19–25	164 (63.8)	93 (36.2)	
• >25	176 (55.7)	140 (44.3)	
comorbidities	296 (61.0)	189 (39.0)	0.381
diabetes	28 (54.9)	23 (45.1)	0.426
coronary disease	107 (60.1)	71 (39.9)	0.996
hypertension	125 (53.6)	108 (46.4)	0.010
peptic ulcer	93 (64.1)	52 (35.9)	0.260
anemia	304 (58.6)	215 (41.4)	0.069

EA – epidural analgesia; IQR – interquartile range

The data of 617 patients diagnosed with gastric cancer that underwent resection of the stomach was retrospectively reviewed and analyzed by univariable and multivariable methods. Patients were divided into two study groups based on the use of epidural analgesia and other methods. The group of patients treated without EA included patients who underwent treatment during a period of time when epidural catheterization use was not the treatment of choice (until the end of 2006); these cases primarily occurred historically earlier than those who were treated with EA. Another reason for non EA administration was the lack of patient consent. Our study included 413 males (66.9%) and 204 females (33.1%) with a median age of 63 (53–71). Epidural analgesia was administered in 246 patients (39.8%). The patients' demographic and clinical characteristics are illustrated in table I and table II.

Analysis of the two group of patients indicated differences in the location of the gastric tumor, the extent of the gastric resection, and spleen removal. Patients treated without EA more frequently experienced malnutrition (BMI < 19). We did not observe statistically significant differences among both studied groups in respect to demographic characteristics, pre-operative risk factors (excluding hypertension), and the length of the procedure. The statistical univariable and multivariable analysis of the factors contributing to postsurgical complications included:

- administered EA,
- gender,
- age,
- pre-surgical BMI,
- diabetes,
- hypertension,

**Table II.** Type and extent of surgical intervention

	Epidural analgesia (EA)		p value
	No n = 371 (%)	Yes n = 371 (%)	
operative approach:			
• laparotomy	270 (72.7)	164 (66.7)	0.104
• thoracolaparotomy	101 (27.3)	82 (33.3)	
type of surgery:			
• gastrectomy (TG)	216 (58.3)	105 (42.7)	<0.001
• TG + distal esophagostomy	97 (26.1)	77 (31.3)	
• distal resection (SG)	55 (14.8)	55 (22.4)	
• proximal gastrectomy (PG)	2 (0.5)	4 (1.6)	
• antrectomy	1 (0.2)	5 (2.0)	
length of surgery (min.):			
• <140	63 (16.9)	62 (25.2)	0.102
• 140–169	93 (25.2)	55 (22.4)	
• 170–209	101 (27.2)	61 (24.8)	
• 210–570	114 (30.7)	68 (27.6)	
perioperative blood transfusion	147 (39.6)	72 (29.3)	0.008
neighboring organ resection	178 (48.0)	110 (44.7)	0.426
splenectomy	138 (37.2)	65 (26.4)	0.005
distal pancreatectomy	15 (4.0)	6 (2.4)	0.282
large bowel resection	6 (1.6)	7 (2.8)	0.298
cholecystectomy	29 (7.8)	14 (5.7)	0.310

- coronary disease,
- and peptic ulcers.

Additionally, in our analysis we included perioperative transfusions, the length of the surgery, and the extent of the multi-organ resection.

### Operative treatment

All cases included in this study contained patients who were operated on by the same experienced (over 30 operations per surgeon) surgical team. Post-operative care and management was provided using consistent post-surgical protocols that included enteral and parenteral nutrition for a period of 7 to 10 days. Total gastrectomy (TG) was performed on 321 patients (52%), 174 (28.2%) patients were treated with TG expanded by resection of the lower section of the esophagus, 110 (17.85%) patients underwent distal subtotal gastrectomy (SG), and in 6 (1%) of these cases proximal gastrectomy (PG) was performed. In 434 (70.3%) of these cases, surgery was performed by laparotomy, in 183 (29.7%) patient's a laparotomy was performed *via* the thoraco-abdominal approach. We performed curative gastrectomies and dissections of the lymph nodes expanded by removal of the additional organs in cases rendering more extensive surgical interventions. The range of surgical resections in both groups of patients is presented in table II.

### Postoperative pain management

All patients (from 2007) were preoperatively evaluated for the postoperative use of epidural analgesia. Additionally, patients

that were administered epidural analgesia consented to the procedure in a separate preoperative assessment. We administered EA in all suitable cases, except in patients with clinical contraindications to the procedure, and in cases where the patient did not consent. Contraindications included:

- coagulation disorders or perioperative use of blood clotting medications,
- inflammation at the catheter placement area,
- neurological conditions.

Prior to administering general anesthesia, in the operating room, the epidural catheter was placed into the epidural space between Th6 and Th7 (when the patient's anatomy dictated, exact vertebral space varied by one up/down segments). The area designated for catheter placement was prepared according to surgical protocols, with the insertion site disinfected and surgical dressing administered. The skin and the subcutaneous tissue in the puncture site was anesthetized with a 2% solution of lidocaine and kept sterile. After the catheter was inserted into the epidural space, it was secured on the skin surface with clearly marked transparent dressing tape. Our postoperative pain management regimen of choice was epidural analgesia, administered *via* continuous infusion of Breivik's mixture into the epidural space using a syringe pump [17]. The mixture was composed of low concentrations of medications (22 µg/ml adrenaline, 2 µg/ml fentanyl and 1.25 mg/ml bupivacaine – which deviates slightly from the standard regimen) in a 0.9% solution of sodium chloride. The epidural infusion delivery rate was about 3:9 ml/h. Patients in both studied groups were intravenously administered coanal-

gesics (metamizol and paracetamol). Patients that were not postoperatively administered EA received a subcutaneously delivered morphine sulfate in fractionated doses (5–10 mg/dose) in 4–6 hour intervals, accompanied with coanalgesics.

### Statistical analysis

Information collected throughout our research was recorded, analyzed, and presented in tables with a cross-tabulation of data. The operative time and age are divided into four categories based on quartiles. The Chi<sup>2</sup> test and Wilcoxon test were used to compare the groups. The relationship between postoperative complications (outcomes) and the use of epidural analgesia is analyzed in a univariable logistic regression model and in a multivariable logistic regression model that controls for confounders. Multi-step forward regression was used to select significant disturbing variables in multivariate models, including significant variables at <0.1 (the multiple variables describing the EA was included in each model regardless of its significance level). The results of the models are presented in the form of odds ratios (OR) and 95% confidence intervals (CI). Variables for which  $p < 0.05$  were considered significant. This analysis is performed with Stata software, version 13.1 (Stata Corporation, College Station, Texas, USA).

### Bioethics

The study was conducted in compliance with the Declaration of Helsinki for medical research and was approved by the Local Bioethics Committee at The Maria Skłodowska-Curie Institute, Oncology Center in Warsaw (permit No. 20/2017 from 09.02.2017). As a retrospective study, according to the approval of the bioethical committee, the informed consent of the patient was not required.

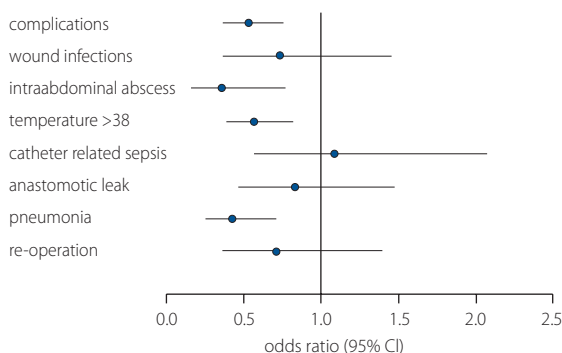
### Results

There were no EA-related complications (neurological deficits, postdural puncture headache), although not every patient managed to insert an epidural catheter. Patients who did not have an epidural catheter inserted for technical reasons were analyzed in the group without EA. Administration of epidural catheters or epidural analgesia is not associated with increased risk for postoperative complications. The postoperative mortality rate was 1%, (6 patients of 617). No thromboembolic or pulmonary complications were present amongst postoperative patients who had received EA. Due to an insignificant occurrence rate, we did not review incidents of hemoperitoneum (intra-abdominal leak), postoperative eventration, or cases of anastomotic strictures (tab. III).

Additionally, the univariable analysis of patients that were administered EA displayed a lower frequency of postoperative complications compared to the group treated with other methods (OR 0.47, 95% CI: 0.34–0.66,  $p < 0.001$ ), intra-abdominal abscesses (OR 0.28, 95% CI: 0.14–0.59,  $p = 0.001$ ), pneumonia (OR 0.39, 95% CI: 0.24–0.63,  $p < 0.001$ ), temperature >38°C (OR

**Table III.** Postoperative complications

Complication type	Epidural analgesia (EA)	
	No n = 371 (%)	Yes n = 246 (%)
overall complications (total)	201 (54.2)	88 (35.8)
temperature >38°C	158 (42.6)	69 (28)
pneumonia	81 (21.8)	24 (9.8)
intra-abdominal abscess	44 (11.9)	9 (3.7)
anastomotic leak	40 (10.8)	20 (8.1)
re-operation	38 (10.2)	14 (5.7)
wound infections	28 (7.5)	13 (5.3)
catheter related sepsis	28 (7.5)	17 (6.9)
anatomic stricture	6 (1.6)	1 (0.4)
intra-abdominal bleeding	4 (1.1)	4 (1.6)
eventration	2 (0.5)	1 (0.4)



**Figure 2.** Risk of complications according to EA administration

0.53, 95% CI: 0.37–0.74,  $p < 0.001$ ) and reoperations (OR 0.53, 95% CI: 0.28–1.00,  $p = 0.049$ ) (fig. 2).

These relationships were confirmed in a multivariable analysis for the general number of complications (OR 0.53, 95% CI: 0.37–0.75,  $p < 0.001$ ), intra-abdominal abscess (OR 0.36, 95% CI: 0.16–0.77,  $p = 0.009$ ), temperature >38°C (OR 0.56, 95% CI: 0.39–0.82,  $p = 0.009$ ), pneumonia (OR 0.42, 95% CI: 0.25–0.71,  $p = 0.001$ ) – tables: IV, V and VI. The relationship between administering EA and reoperation in a multivariable analysis was not confirmed. Diagnosis of pneumonia was based on the correlation of clinical symptoms and radiological determinations. There were no significant statistical differences in univariable and multivariable analysis of wound infections, infection of the central line, or the anastomotic stricture, (table VI and table VII).

### Discussion

Complications associated with gastrectomy with D2 lymphadenectomy for the treatment of gastric cancer presents multiple clinical considerations against extensive lymphadenectomy [10, 13, 14]. The overall rate of complications is between 17 to 48%. The most frequent postoperative complications in gastric resection surgeries for curative gastric

**Table IV.** Analysis of complications (total complications and temperature >38°C) based on the administration of epidural analgesia

variable	Temperature >38°C											
	Total complications						Temperature >38°C					
	Univariable			Multivariable			Univariable			Multivariable		
	OR	95%CI	p value	OR	95%CI	p value	OR	95%CI	p value	OR	95%CI	p value
EA	0.47	0.34–0.66	<0.001	0.53	0.37–0.75	<0.001	0.53	0.37–0.74	<0.001	0.56	0.39–0.82	0.003
BMI:												
• 19–25	1.49	0.77–2.88	0.241	1.67	0.81–3.44	0.162	1.41	0.69–2.88	0.341	1.61	0.75–3.45	0.225
• >25	1.68	0.88–3.24	0.117	2.25	1.09–4.65	0.029	1.77	0.88–3.56	0.111	2.38	1.11–5.10	0.026
diabetes	1.31	0.74–2.32	0.363				1.59	0.89–2.83	0.115			
CD	1.16	0.82–1.64	0.410				1.02	0.71–1.46	0.925			
hypertension	0.67	0.49–0.94	0.019	0.64	0.44–0.94	0.023	0.75	0.53–1.05	0.094			
peptic ulcer	1.29	0.89–1.88	0.178				1.68	1.15–2.44	0.007	1.88	1.25–2.81	0.002
OT (ref. <140):												
• 140–169	1.81	1.10–3.00	0.021	1.43	0.83–2.47	0.194	1.82	1.06–3.13	0.030			
• 170–209	2.38	1.45–3.89	0.001	1.58	0.92–2.69	0.096	2.51	1.48–4.23	0.001			
• 210–570	3.39	2.09–5.51	0.000	2.18	1.28–3.70	0.004	2.72	1.63–4.54	<0.001			
BT	1.69	1.21–2.36	0.002	1.58	1.10–2.28	0.014	1.50	1.07–2.10	0.019	1.45	1.00–2.10	0.047
Spl	2.77	1.95–3.92	<0.001	2.17	1.49–3.18	<0.001	2.73	1.93–3.87	<0.001	2.57	1.77–3.72	0.000
SP	2.33	0.93–5.87	0.071				1.59	0.66–3.80	0.299			
gender (male)	1.85	1.31v2.61	<0.001	1.60	1.09–2.33	0.016	1.63	1.14–2.33	0.008	1.59	1.08–2.35	0.020
age (ref. <53):												
• 54–62	1.23	0.78–1.95	0.367	1.39	0.84–2.32	0.203	1.02	0.64–1.64	0.921	0.96	0.58–1.61	0.885
• 63–70	1.89	1.18–3.03	0.008	2.05	1.21–3.48	0.007	1.60	0.99–2.58	0.053	1.49	0.89–2.49	0.131
• 71–87	1.00	0.64–1.58	0.995	1.21	0.71–2.05	0.483	0.81	0.50–1.30	0.385	0.75	0.45–1.28	0.293

EA – epidural analgesia; CD – coronary disease; BT – blood transfusion; OT – operation time (minutes); Spl – splenectomy; SP – splenopancreatectomy

**Table V.** Analysis of complications (pneumonia and intra-abdominal abscess) based on the administration of epidural analgesia

Variable	Pneumonia						Intra-abdominal abscess					
	Univariable			Multivariable			Univariable			Multivariable		
	OR	95%CI	p value	OR	95% CI	p value	OR	95% CI	p value	OR	95% CI	p value
EA	0.39	0.24–0.63	<0.001	0.42	0.25–0.71	0.001	0.28	0.14–0.59	0.001	0.36	0.16–0.77	0.009
BMI:												
• 19–25	1.03	0.41–2.62	0.948	1.28	0.48–3.46	0.621	1.28	0.37–4.47	0.699			
• >25	1.58	0.64–3.90	0.323	2.74	1.03–7.30	0.044	1.33	0.39–4.57	0.652			
diabetes	1.56	0.79–3.10	0.199	1.91	0.90–4.03	0.092	0.90	0.31–2.60	0.843			
CD	1.16	0.74–1.83	0.522				0.97	0.52–1.81	0.927			
hypertension	0.64	0.41–1.01	0.057	0.58	0.35–0.96	0.036	0.57	0.30–1.07	0.078			
peptic ulcer	1.64	1.03–2.60	0.037	1.82	1.11–3.01	0.018	1.19	0.62–2.25	0.601			
OT (ref. <140):												
• 140–169	1.59	0.16–16.02	0.694				2.91	0.92–9.17	0.068			
• 170–209	1.86	0.21–16.47	0.578				2.42	0.76–7.69	0.134			
• 210–570	2.27	0.27–19.38	0.453				4.59	1.55–13.59	0.006			
BT	2.30	1.16–4.55	0.017	2.34	1.13–4.85	0.023	2.82	1.59–5.00	<0.001	2.55	1.37–4.73	0.003
Spl	2.27	1.04–4.92	0.038	2.10	0.93–4.74	0.075	6.79	3.59–12.85	<0.001	5.61	2.91–10.82	<0.001
SP	1.68	0.55–5.12	0.363				4.67	1.73–12.61	0.002			
gender (male)	2.56	1.51–4.34	<0.001	2.39	1.37–4.17	0.002	3.00	1.38–6.48	0.005	2.93	1.27–6.75	0.012
age (ref. <53):												
• 54–62	0.68	0.37–1.27	0.228	0.68	0.37–1.27	0.228	0.99	0.45–2.16	0.978			
• 63–70	1.08	0.60–1.94	0.804	1.08	0.60–1.94	0.804	1.11	0.51–2.43	0.794			
• 71–87	0.86	0.48–1.54	0.607	0.86	0.48–1.54	0.607	0.61	0.26–1.43	0.252			

EA – epidural analgesia; CD – coronary disease; BT – blond transfusion; OT – operation time (minutes); Spl – splenectomy; SP – splenopancreatectomy

**Table VI.** Analysis of complications (anastomotic leak and reoperation) based on administration of epidural analgesia

Variable	Anastomotic leak						Re-operation					
	Univariable			Multivariable			Univariable			Multivariable		
	OR	95% CI	p value	OR	95% CI	p value	OR	95% CI	p value	OR	95% CI	p value
EA	0.73	0.42–1.29	0.278	0.83	0.46–1.48	0.523	0.53	0.28–1.00	0.049	0.72	0.37–1.40	0.328
BMI:												
• 19–25	0.53	0.20–1.42	0.208				0.62	0.22–1.76	0.373			
• >25	0.76	0.30–1.94	0.570				0.76	0.28–2.08	0.591			
diabetes	2.16	1.00–4.70	0.051	2.35	1.07–5.19	0.034	1.20	0.45–3.17	0.712			
CD	1.06	0.59–1.90	0.836				1.11	0.60–2.05	0.750			
hypertension	0.81	0.46–1.42	0.457				0.78	0.43–1.44	0.431			
peptic ulcer	1.59	0.89–2.83	0.119				0.76	0.37–1.55	0.449			
OT (ref. <140):												
• 140–169	1.09	0.39–3.02	0.866				0.96	0.34–2.73	0.944			
• 170–209	2.37	0.97–5.81	0.058				1.59	0.62–4.08	0.330			
• 210–570	2.56	1.07–6.14	0.035				2.44	1.01–5.87	0.047			
BT	2.10	1.23–3.59	0.007	2.06	1.19–3.56	0.009	4.70	2.54–8.68	<0.001	4.77	2.53–9.01	<0.001
Spl	1.91	1.12–3.27	0.018	1.78	1.03–3.09	0.039	3.37	1.88–6.04	<0.001	2.90	1.58–5.33	0.001
SP	0.98	0.22–4.30	0.975				3.65	1.28–10.40	0.015			
gender (male)	1.70	0.91–3.17	0.095				3.44	1.52–7.77	0.003	3.51	1.51–8.14	0.003
age (ref. <53):												
• 54–62	0.92	0.42–2.02	0.830				1.02	0.44–2.34	0.972			
• 63–70	1.36	0.64–2.90	0.421				1.64	0.74–3.60	0.221			
• 71–87	0.94	0.43–2.04	0.872				0.73	0.30–1.77	0.482			

EA – epidural analgesia; CD – coronary disease; BT – blond transfusion; OT – operation time (minutes); Spl – splenectomy; SP – splenopancreatectomy

**Table VII.** Analysis of complications (wound infections and catheter related sepsis) based on the administration of epidural analgesia

Variable	Wound Infections				Catheter related sepsis				
	OR	95% CI	p value	OR	95% CI	p value	OR	95% CI	p value
EA	0.68	0.35–1.35	0.272	0.73	0.37–1.45	0.368	0.91	0.49–1.70	0.766
BMI:									
• 19–25	3.05	0.39–23.49	0.285	0.79	0.22–2.86	0.716	0.79	0.22–2.86	0.716
• >25	3.38	0.44–25.64	0.240	1.33	0.39–4.57	0.652	1.33	0.39–4.57	0.652
diabetes	0.87	0.26–2.92	0.820	2.21	0.93–5.24	0.072	2.21	0.93–5.24	0.072
CD	1.02	0.51–2.05	0.951	1.72	0.92–3.20	0.089	1.78	0.92–3.42	0.085
hypertension	0.58	0.29–1.19	0.139	1.11	0.60–2.06	0.748	1.11	0.60–2.06	0.748
peptic ulcer	0.91	0.42–1.95	0.809	1.52	0.79–2.94	0.214	1.52	0.79–2.94	0.214
OT (ref. <140):									
• 140–169	1.50	0.43–5.25	0.524	1.50	0.43–5.25	0.524	1.50	0.43–5.25	0.524
• 170–209	2.64	0.84–8.30	0.097	3.55	1.16–10.82	0.026	3.55	1.16–10.82	0.026
• 210–570	3.12	1.02–9.50	0.046	3.12	1.02–9.50	0.046	3.12	1.02–9.50	0.046
BT	1.62	0.86–3.07	0.136	1.50	0.81–2.77	0.195	1.50	0.81–2.77	0.195
Spl	1.65	0.87–3.14	0.124	2.28	1.24–4.19	0.008	2.52	1.32–4.80	0.005
SP	3.55	1.14–11.10	0.029	2.85	0.90–9.02	0.075	0.63	0.08–4.78	0.653
gender (male)	2.52	1.10–5.80	0.029	2.38	1.03–5.50	0.042	0.54	0.29–0.99	0.047
age (ref. <53):									
• 54–62	1.08	0.41–2.81	0.879	0.96	0.36–2.57	0.940	0.98	0.36–2.66	0.961
• 63–70	1.74	0.70–4.28	0.230	2.45	1.04–5.80	0.041	2.32	0.96–5.60	0.062
• 71–87	0.92	0.34–2.44	0.860	0.92	0.34–2.44	0.860	0.94	0.34–2.61	0.902

EA – epidural analgesia; CD – coronary disease; BT – blond transfusion; OT – operation time (minutes); Spl – splenectomy; SP – splenopancreatectomy



cancer interventions are pneumonia, surgical site infections, (incision infections, intra-abdominal abscesses) and leaking anastomosis [9–14, 18]. Despite the potential for postoperative complications, extensive surgical resection with lymph nodes dissection remains the only curative therapy for gastric cancer worldwide. Experienced medical institutions specializing in surgical oncology routinely perform extensive curative resections for gastric cancer [9, 10, 14, 19].

Effective analgesia is an essential part of postsurgical management and provides statistically and clinically significant improvements in treatment outcomes. Most published clinical studies have demonstrated that the administration of epidural analgesia in gastric surgery patients is a safe practice as a means to improve perioperative outcomes [16, 20–24]. Effective postoperative pain management, as well as the reduction of stress response to surgery along with management of the cardiovascular system and microcirculation significantly reduces complications. Furthermore, studies suggest that administering EA contributes to the reduction of perioperative blood loss. The recommended technique requires continuous infusion of pain medications assisted by intermittent bolus injections [25, 26].

A comprehensive literature review of the effect of postoperative analgesia on surgical outcomes [24] showed the impact of administering epidural analgesia on complications rates following major abdominal surgery. Throughout this study, authors established that the administration of epidural analgesia significantly reduces the risk of pulmonary and cardiovascular complications, as well as thromboembolism, postoperative occlusions, and hastens the return of bowel function. Our study was performed retrospectively and is therefore subject to associated biases. During the extensive research period in which the review of this data occurred, our standards of postoperative care and surgical experience have improved; possibly affecting our findings had this data included newer cases. Therefore, based on this study alone, we cannot definitively conclude that administering EA decreases the risk of complications after gastrectomy. There is, however increasing evidence of the overall positive impact that EA has on treatment outcomes. When considering the retrospective review of 84 patients that underwent laparoscopic SG [27], all data suggests that administering EA has no significant impact on treatment outcomes, except for patients treated with EA who experienced urinary retention.

In a prospective study of 1021 patients, the analysis confirmed more effective pain management, a lower need for analgesics, and a shorter stay in the intensive care unit [28]. No statistically significant differences were reported for mortality and the postoperative complication rate. Further analysis demonstrated a reduction of postoperative complications in the group of patients administered EA that underwent vascular surgical interventions. In the relatively smaller groups of patients that underwent gastrectomy (77 patients), large

intestine or bile duct operation, the difference between the number of postoperative complications remains insignificant. The results of the Cochrane Database analysis [29] in which 94 studies were evaluated, (total of 5864 patients) suggests effective pain management and an accelerated return of gastrointestinal transit in patients treated with EA. With the use of the open surgery technique, EA reduces the length of the hospital stay. There was no difference in vomiting incidence or anastomotic leak. Complications of epidural analgesia are rare, but additional studies to examine the impact of administering epidural analgesia in extensive surgical interventions for gastric cancer are needed.

A recently published retrospective review of the American College of Surgeons National Surgical Quality Improvement Program [30] performed for patients undergoing open elective esophagectomies and gastrectomies for nonmetastatic cancer, analyzed a group of 2599 gastrectomies, among which 18% received EA. The only conclusion from the analysis is that EA was associated with a longer length of stay (EA median [IQR] 8 [7, 11] vs. no EA 7 [6, 11],  $p = 0.0002$ ). No other differences between the groups were noted.

Of the retrospective review of 723 gastric cancer resections performed at our institution, 617 cases had complete medical documentation that was adequate for review (85.3%). Data not included in this study amounting to the remaining 14.7% of cases was excluded due to random issues such as incomplete medical records and other associated factors. The analyzed group of patients was treated with comparable surgical techniques, postoperative care, and perioperative management protocols. Patients administered EA did not experience a higher number of complications than the group of patients treated with other methods. Thus, administering EA has proven to be safe in the perioperative care of patients undergoing gastric resection. Research indicates that the frequency of wound infections (fever  $> 38^{\circ}\text{C}$ , intra-abdominal abscess) pneumonia and reoperations is reduced in the group of patients with EA. Metaanalysis [26] as well as our assessments confirm that effective postoperative pain management decreases the incidence of pulmonary complications. We observed a decreased number of other complications, (except for frequency of anastomotic leak), however in conclusion, they offer no statistical significance.

## Conclusions

Our findings indicate that administering EA to patients undergoing major stomach resection for gastric cancer is safe. Furthermore, postoperative treatment with epidural analgesia following stomach resection contributes to a reduction in the number of postoperative complications; this is most notable in the reduced number of cases of pneumonia, sepsis, and the need for additional surgical interventions.

**Conflict of interest:** none declared

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Received: 20 Mar 2021

Accepted: 23 May 2021

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