

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 approaches and improvements in surgical techniques. The European Society for Medical Oncology (ESMO) guidelines of treatment for patients diagnosed with an advanced GC include perio-

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perative 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).

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

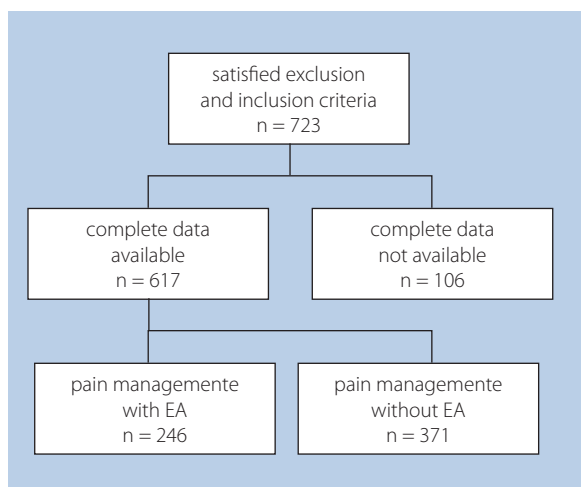


Figure 1. Flow-chart of the study

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 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,
- 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.

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 |
| • thoracolumbarotomy | 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 |

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 coanalgesics (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 Chi² 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 $>38^{\circ}\text{C}$ (OR 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%

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 $>38^{\circ}\text{C}$ | 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) |

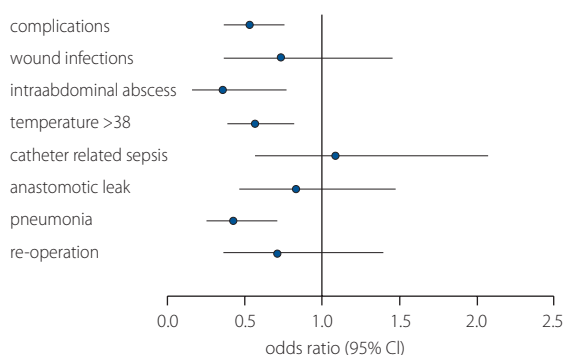


Figure 2. Risk of complications according to EA administration

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, $p = 0.001$) – 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 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

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 | 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 | Multivariable | OR | 95% CI | p value | Multivariable | | | | |
| 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 | 1.08 | 0.57–2.08 | 0.808 |
| BMI: | | | | | | | | | | | | |
| • 19–25 | 3.05 | 0.39–23.49 | 0.285 | | | | 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 | | | |
| diabetes | 0.87 | 0.26–2.92 | 0.820 | | | | 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 | | | |
| peptic ulcer | 0.91 | 0.42–1.95 | 0.809 | | | | 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 | | | |
| • 170–209 | 2.64 | 0.84–8.30 | 0.097 | | | | 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 | | | |
| BT | 1.62 | 0.86–3.07 | 0.136 | | | | 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 | 0.50 | 0.26–0.94 | 0.032 |
| 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

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.

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