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1	Preoperative Embolization of the Splenic and Left Gastric
2	Arteries Does Not Seem to Decrease the Rate of Anastomotic
3	Leaks after Esophagogastroctomy
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### 9 Abstract

<sup>10</sup> Introduction: Embolization of the gastric and splenic artery is performed to improve blood

<sup>11</sup> flow to the constructed gastric tube during esophagectomy and thus reduce the rate of

<sup>12</sup> anastomotic leaks for this type of surgery.Patients and Methods: This is a retrospective,

<sup>13</sup> comparative study conducted from 2003 to 2013 of 32 patients undergoing preoperative

<sup>14</sup> embolization versus 102 patients not undergoing embolization to assess the impact of this

15 technique on the rate of anastomotic leakage. The variables considered were age, gender,

<sup>16</sup> comorbidities, preoperative histopathology, location of the tumor, type of neoadjuvant therapy

17 (if appropriate), type of surgery and type of anastomosis.

18

#### 19 Index terms—

### <sup>20</sup> 1 I. Introduction

nastomotic leakage after esophagectomy is a complication varying in severity that may increase morbidity and
mortality in patients depending on the location of the anastomosis (thoracic or cervical). The incidence of
anastomotic leaks ranges from 3% to 25%, especially for cervical esophagogastric anastomoses 1,2,3,4.

Esophageal anastomotic leakage is defined as the extravasation of gastric/esophageal luminal contents. Bacterial contamination may cause local abscess, fistula, anastomotic dehiscence, surgical wound dehiscence, sepsis and even death.

Esophagogastric anastomosis may be caused by patient's status (malnutrition, immunosuppression...) and/or the surgical technique employed (insufficient perfusion of the constructed gastric tube, location of the anastomosis,

29 tense anastomosis...) 5.

Embolization of the left gastric artery before esophagectomy has been reported to improve ischemia in the constructed gastric tube during anastomosis.

The objective of this study was to assess the rate of anastomotic leakage in patients undergoing embolization prior to esophagectomy versus that of patients not undergoing preoperative embolization.

# <sup>34</sup> 2 II. Patients and Methods

<sup>35</sup> This is a retrospective, observational study of a cohort of patients undergoing esophagectomy either for a malign

or benign condition referred to the Esophagogastric Surgery Unit of a third-level hospital between June 2003 and December 2013. All patients were examined preoperatively by a multidisciplinary board. Data were prospectively

38 collected in a database.

39 We compared the rate of anastomotic leaks in patients undergoing embolization of the left gastric A Water-

40 soluble contrast studies were performed on all patients for anastomotic leakage on postoperative day 7 before 41 oral feeding was initiated. Anastomotic leakage can be detected on X-ray scans and manifest either clinically or biologically in blood tests. When a leak was detected, we used a classification system based on clinical, radiological and endoscopic findings. Then, the protocol established and previously published for the management of anastomotic leaks was implemented. 6 (Figure 2). artery and splenic artery before esophagectomy with that of patients not undergoing preoperative embolization. The following variables were retrospectively analyzed: age, gender, comorbidities, tumor location, administration and type of neoadjuvant treatment, type of surgery, histology of the resected tissue, postoperative complications including type of anastomotic leakage and management of the leak.

Embolization was performed by interventional radiologists three weeks before surgery. Under local anesthesia, the femoral right artery was punctured. Embolization of the splenic artery was performed using a 8mm-Amplatzer®, whereas the left gastric artery was embolized using coils (Figure 1).

# 52 **3** IV. Results

A total of 138 patients with esophageal cancer and 4 patients with benign lesions requiring esophagectomy were recorded in the database during the study period. Of the 138 patients registered, 121 were male and 19 were female, with an average age of 60 years.

Figure 3 shows the process of inclusion and exclusion for embolization candidates. Of a total of 142 patients, 56 108 did not undergo preoperative embolization, whereas 34 underwent embolization of the left gastric artery 57 and splenic artery before esophagectomy. Of the patients undergoing embolization, two underwent coloplasty, in 58 one due to the absence of the right gastro-omental artery and in the other case because his tumor at the distal 59 esophagous extended more than 7cm to the stomach and the gastric tube could not be constructed with sufficient 60 tumor-free margins. Finally, a total of 32 patients undergoing embolization were included in the study. As to the 61 108 patients without embolization, one was candidate to coloplasty for esophageal necrosis and partial gastric 62 necrosis caused by caustic ingestion. In other case, the patient underwent endoscopic resection because the tumor 63 was located at the mucous membrane and the patient presented substantial comorbidities. Four patients had 64 an unresectable tumor. Finally, 102 patients undergoing esophagectomy without preoperative embolization and 65 developping esophageal and gastric anastomosis were included in the study. 66 Therefore, of the 134 candidates to esophagectomy, 32 (23.8%) underwent embolization and 102 (76.2%) had 67

no embolization. Table I shows demographics and comorbidities. No statistically significant differences were 68 found between patients undergoing embolization and patients not undergoing embolization. Table II shows the 69 histopathological findings following esophagectomy and the location of the esophageal tumors. No statistically 70 significant differences were found between patients undergoing embolization and patients not having embolization. 71 72 Table III shows the preoperative treatment administered and the type of surgery performed. Of the 32 patients 73 undergoing embolization, 28 (87.5%) received chemoradiotherapy preoperatively, whereas 52 of the 102 (50.9%) 74 patients without embolization received chemoradiotherapy preoperatively (p=0.0001). Statistically significant 75 differences were found between the embolization group and the no-embolization group (p=0.0001). As regards the type of surgery, no statistically significant differences were found between patients undergoing embolization 76 and patients not undergoing embolization. Minimally invasive esophagectomy was performed in 30 (93.75%) of 77 the patients with embolization and in 93 (91.17%) of the patients without embolization. As to the location of 78 the anastomosis (cervical vs thoracic), no statistically significant differences were observed (p=0.586). 79

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As many as 14 patients (43.75%) undergoing embolization developed anastomotic leaks, of which 6 (18.75%) 82 were detected on X-ray scans or due to mild symptomatology. Eight leaks manifested clinically. As to the patients 83 without embolization, 33 (32.35%) developped anastomotic leakage, of which 12 (11.7%) had clinical symptoms. 84 As regards the development of an astomotic leakage, no statistically significant differences were found between the 85 two groups No statistically significant differences were found between the two groups concerning the incidence of 86 the most severe anastomotic leaks (type III and IV) (p=0.087). (See Table IV). No complications were associated 87 with embolization. Regarding long-term complications, two (6.25%) of the patients with embolization and four 88 (3.92%) of the patients without embolization developed anastomotic stricture formation. Three (9.37%) patients 89 with embolization died soon after surgery (one died of pneumony and two of anastomotic leakage); ten (9.8%)90 patients without embolization died (four died of anastomotic leakage, one of liver failure caused by existing 91 cirrhosis, one of heart failure and four of respiratory complications). (See Table IV). 92

### 93 5 V. Discussion

94 This non-randomized retrospective study revealed that no statistically significant differences exist concerning 95 anastomotic leak rates between patients undergoing embolization and patients not undergoing embolization

96 before esophagectomy.

Anastomotic leakage is a severe complication of esophageal surgery. The incidence of this complication ranges between 10 and 25% 7,8 and may increase mortality rates up to 50% 9. Anastomotic leakage is defined as a leak of luminal contents from a surgical join between two hollow viscera 10. The escape of luminal contents may

100 cause local abscess, fistula, sepsis or death.

Variations in reported rates of esophageal anastomotic leaks may be due to the different definitions used to 101 classify this complication and its location; this also has an impact on therapeutic algorithms 7,11,12. With 102 regard to the physiopathogeny of anastomotic esophagogastric dehiscences Turkyilmaz et al. 7 defined four types 103 according to the triggering factors; thus dehiscences may be caused : a) by a systemic disease; b) by the intrinsic 104 anatomy of the esophagus; c) by the technical factors of the surgery; d) by postoperative care and complications. 105 As to the classification of esophagogastric anastomotic fistulae, like Lerut el al 11, we classify them according 106 to their clinical impact as assessed through clinical, radiological and endoscopic examination and to the therapy 107 required 6. The gastric flow has been proven to decrease both in animals and humans by more than 70% after 108 resection of the left gastric artery, the short vessels and their veins, which is required to rise the constructed 109 gastric tube 13,14,15. The decrease in gastric flow results in ischemia. 110

## 111 6 Volume

Different authors have described a variety of techniques for improving blood flow and venous drainage in esophagogastric anastomosis and reduce the rate of anastomotic leakage and its severe effects, with different outcomes 16,17,18.

To improve microvascularization in the reconstructed gastric tube through ischemic conditioning of the 115 stomach, Akiyama et al 19 described the technique of embolization of the left gastric artery and splenic artery 116 through the femoral artery 12 days prior to the esophagectomy. Although Akiyama did not report statistically 117 significant differences in anastomotic leak rates, he demonstrated that vascularization of the reconstructed gastric 118 tube at the moment of esophagogastric anastomosis improved in patients undergoing embolization. Diana et 119 al 20 did not found statistically significant differences either between patients with embolization and patients 120 without embolization. However, they observed that patients undergoing embolization were more likely to develop 121 esophagogastric stricture (32% vs 3%). 122

Similarly, transient bloodletting of the short gastric vein seems to be an efficient mechanism for conditioning microcirculation, which improves the circulation of the oral side of the gastric tube during esophagectomy 17. However, outcomes have not been reported.

A different method to improve microcirculation is performing delayed conditioning by two-stage esophageal surgery 21,22,23 (the gastric tube is reconstructed in the first stage and esophagectomy is performed some days later in a second stage).

Yoshimi et al 16 revascularized the gastric tube using the splenic artery and vein, external carotid artery, and internal jugular vein in 21 patients who did not develop anastomotic leakage. However, differences in anastomotic leak rates between this group and the control group were not statistically significant.

This study has several limitations. Firstly, this is not a randomized, prospective, case control study. Secondly, the sample of patients undergoing embolization was very small, which reduced its statistical power. Third, chemoradiotherapy was administered to the two groups differently, which may affect outcomes regarding the rate of anastomotic leakage.

In conclusion, according to the results of this study, it cannot be concluded that embolization of the left gastric artery and the splenic artery before esophagectomy reduces the rate of anastomotic leaks.

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Figure 1: Figure 1 :



Figure 2: Figure 2 :



Figure 3: Figure 3 :



Figure 4: 3

Ι

	WITH EMBOLIZA- TION (n=32)	WITHOUT	p-value
		EMBOLIZATION	
		(n=102)	
AGE (YEARS)	61 (46-75)	59.95 (29-88)	0.603
Gender			
Female	5~(15%)	14 (13.7%)	0.982
Male	27~(85%)	88~(86.2%)	
Comorbidities			
Smoker	13~(40%)	37~(36.3%)	0.679
Drinker	0	13(12.6%)	0.065
Heart Disease	13~(40.6%)	39~(38.23%)	0.837
COPD	6~(18.7%)	14 (13.7%)	0.570
Obesity	2~(6.2%)	1 (0.98%)	0.141
Diabetes	1 (3.1%)	9~(8.8%)	0.450
Chronic liver disease	1 (3.1%)	4(3.9%)	1.000

Figure 5: Table I :

# $\mathbf{II}$

	WITH	WITHOUT	p-
	EMBOLIZAT	IONEMBOLIZATION	value
	(n=32)	(n=102)	
HISTOPATHOLOGY			
Adenocarcinoma	16~(50%)	48~(47.05%)	0.684
Ca. Epidermoid	16~(50%)	47(46.07%)	0.676
Ca. Neuroendocrine	0	1 (0.98%)	-
GIST	0	3~(2.94%)	-
Other (achalasia, necrosis, peptic	0	3(2.94%)	-
stricture)			
LOCATION			
Proximal esophagus	9~(28.1%)	21~(20.58%)	0,267
Mediastinal esophagus	14 (13.7%)	60~(58.82%)	
Distal esophagus	9~(28.1%)	18~(17.64%)	

Figure 6: Table II :

 $\mathbf{III}$ 

Valu EMBOLIZATION EMBOLIZATION (n=32) (n=102) PREOPERATIVE CHEMOHERAPY/RADIATION	
PREOPERATIVE CHEMOHERAPY/RADIATION	e
CHEMOHERAPY/RADIATION	
THERAPY	
No 4 (12.5%) 50 (49.1%) 0.00	01
Yes 28 (87.5%) 52 (50.9%)	
SURGERY	
Ivor-Lewis $10 (31.25\%)$ $37 (36.2\%)$ $0.692$	3
Transhiatal esophagectomy $3 (9.37\%)$ $13 (12.74\%)$	
Three-stage esophagectomy $19 (59.37\%) 52 (50.9\%)$	
$ \begin{array}{llllllllllllllllllllllllllllllllllll$	
LOCATION OF THE ANASTO-	
MOSIS	
Cervical $21(65.62\%)$ $65(63.7\%)$ $0.58\%$	6
Thoracic $11(34.38\%)$ $37(36.3\%)$	

Figure 7: Table III :

IV

	WITH	WITHOUT	p-
	EMBOLIZATION	EMBOLIZATION	value
	(n=32)	(n=102)	
ANASTOMOTIC	14 (43.75%)	33~(32.35%)	0.289
LEAKS			
Type I	2(6.25%)	11 (10.78%)	
Type II	4 (12.5%)	10 (9.8%)	
Type III	6 (18.75%)	10 (9.8%)	
Type IV	2(6.25%)	2~(1.9%)	
III-IV leaks	8 (25%)	12 (11.7%)	0.087
Stricture	14(13.7%)	4 (3.92%)	0.628
Deaths ( $<30$ postopera-	3~(9.37%)	10 (9.8%)	1
tive days )			

Figure 8: Table IV :

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