Worldwide practice in gastric cancer surgery

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Abstract

AIM: To evaluate the current status of gastric cancer surgery worldwide.

METHODS: An international cross-sectional survey on gastric cancer surgery was performed amongst international upper gastro-intestinal surgeons. All surgical members of the International Gastric Cancer Association were invited by e-mail to participate. An English web-based survey had to be filled in with regard to their surgical preferences. Questions asked included hospital volume, the use of neoadjuvant chemotherapy, preferred surgical approach, extent of the lymphadenectomy and preferred anastomotic technique. The invitations were sent in September 2013 and the survey was closed in January 2014.

RESULTS: The corresponding specific response rate was 227/615 (37%). The majority of respondents: originated from Asia (54%), performed > 21 gastrectomies per year (79%) and used neoadjuvant chemotherapy (73%). An open surgical procedure was performed by the majority of surgeons for distal gastrectomy for advanced cancer (91%) and total gastrectomy for both early and advanced cancer (52% and 94%). A minimally invasive procedure was preferred for distal gastrectomy for early cancer (65%). In Asia surgeons preferred a minimally invasive procedure for total gastrectomy for early cancer also (63%). A D1+ lymphadenectomy was preferred in early gastric cancer (52% for distal, 54% for total gastrectomy) and a D2 lymphadenectomy was preferred in advanced gastric cancer (93% for distal, 92% for total gastrectomy)

CONCLUSION: Surgical preferences for gastric cancer surgery vary between surgeons worldwide. Although the majority of surgeons use neoadjuvant chemotherapy, minimally invasive techniques are still not widely adapted.

Key words: Gastric cancer; Gastrectomy; Laparoscopy;
Core tip: Since surgical techniques might differ over time and between countries, we aimed to evaluate international preferences in gastric cancer surgery by means of a cross-sectional survey. Surgical preferences for gastric cancer surgery vary between surgeons worldwide. Minimally invasive gastrectomy is still not widely adapted, but most popular in Asia to treat patients with early gastric cancer. Neo-adjuvant chemotherapy is used by the majority of surgeons worldwide. A D1+ lymphadenectomy is preferred for early gastric cancer and a D2 lymphadenectomy is preferred for advanced gastric cancer.

INTRODUCTION

Gastric cancer is the fifth most common type of cancer worldwide. Its treatment consists of (neo-)adjuvant chemotherapy and/or chemoradiation and surgical resection of the tumor and lymph nodes. The worldwide surgical practices may vary between surgeons, countries and continents. Current topics of debate in gastric cancer surgery are: (1) The influence of volume of hospitals and surgeons on the outcome after gastrectomy; (2) The technique of surgery: open or minimally invasive gastrectomy; (3) Reconstruction of the alimentary tract by means of a jejunal pouch. (4) The extent of lymph node dissection and need for omental resection and/or pancreatosplenectomy; and (5) The type of (neo-)adjuvant treatment in patients with gastric cancer.

In this article the current practice of surgeons worldwide will be evaluated by means of a survey.

MATERIALS AND METHODS

An international cross-sectional survey about the surgical treatment of gastric cancer was performed amongst international gastric surgeons. All surgical members of the International Gastric Cancer Association (IGCA) were invited by email to participate after approval of the IGCA was obtained. An English web-based survey had to be filled in according to the surgeons’ preferences. Questions asked included hospital volume, the use of neoadjuvant treatment, preferred surgical approach, extent of the lymphadenectomy and preferred anastomatic technique and are attached to the manuscript (Appendix 1). Definitions of the extent of lymph node dissection and gastric cancer classification were according to the Japanese gastric cancer classification system. The invitations were sent in September 2013 and the survey was closed in January 2014. Statistical analysis was performed with the $\chi^2$ test using the IBM SPSS Statistics (version 21; IBM Corporation, Armonk, NY, United States). Data were considered significant if $P < 0.05$.

RESULTS

Demographics

The survey was completed by 248 of 615 (40%) members of the IGCA. The 21 duplicate respondents were consequently excluded. The corresponding specific response rate was therefore 227/615 (37%). The respondents originated from Asia (54%), Europe (27%), South America (12%), North America (6%), Africa (0.4%), and Oceania (0.4%) (Figure 1).

Volume

The volume of the participating surgeons was $\leq 10$ gastrectomies per year in 16 (7%) respondents and $> 21$ resections in 180 (79%) respondents. In Asia, the majority of respondents (57%) performed $> 61$ resections (Figure 2). Medium and high volume surgeons worked in a university hospital (74%) more often than in a regional hospital (5%, $P = 0.048$).

Open vs minimally invasive gastrectomy

The current survey revealed that minimally invasive distal gastrectomy was preferred by 65% of surgeons in the treatment of early gastric cancer. The Asian respondents performed minimally invasive distal gastrectomy for early gastric cancer in 82% of the cases (Figure 3). In South America minimally invasive and open distal gastrectomy were equally performed. Minimally invasive distal gastrectomy for advanced gastric cancer was performed by only 9% of respondents. These results were comparable in all continents. For total gastrectomy, minimally invasive total techniques were favored by 49% for early gastric cancer and by 6% for advanced gastric cancer. However, in Asia the majority (64%) of respondents performed minimally invasive total gastrectomy for early gastric cancer, whereas other continents preferred the open procedure. For total gastrectomy for advanced cancer, there was no difference between continents.

Anastomoses

The preference of 83% of the participating surgeons in the survey was to construct a direct esophagojejunostomy without jejunal pouch reconstruction after total gastrectomy. In merely 17% of surgeons a pouch was the preferred method of reconstruction. This percentage was consistent between all continents.
A pouch reconstruction was slightly more popular amongst surgeons from a university hospital than surgeons from a regional hospital (19% vs 11%, \( P = 0.50 \)). The data from the survey reveal that anastomoses were preferably performed by means of a mechanical stapler by 92% of respondents, compared to 8% of surgeons who favored a hand-sewn anastomosis.

**Extent of dissection**

The current survey indicated that surgeons preferred a D1+ resection in 52% of distal gastrectomies and 54% of total gastrectomies for early cancers (Figure 4). In Asia and Northern America a D2 dissection was performed less frequently for early stage tumors compared to Europe and Southern America (Table 1). A D2 resection was favored by 93% of distal gastrectomies and 92% of total gastrectomies for advanced tumors. Resection of the spleen was preferably performed by 20% of all respondents: 33% of Asian respondents, 19% of South American respondents, and 15% of European respondents. The survey reveals that resection of the greater omentum was preferred by 89% of the participating surgeons.

**(Neo)adjuvant therapy**

The results of our survey on the use of neoadjuvant treatment for gastric cancer indicated that chemotherapy was preferred by 73% and chemoradiation was favored by 12% of respondents. Only 16% favored treatment without neoadjuvant treatment. These results did not differ significantly over continents (Table 2).

**DISCUSSION**

In this study the current worldwide trends in gastric surgery for cancer were evaluated by means of a survey amongst surgeons. It was found that the majority of surgeons have a high annual volume of gastrectomies. Open gastrectomy was still the preferred procedure in most procedures (68%)
> 20 gastrectomies per year which is considered a high volume in literature. With regard to annual volume of resection, both the volume of the individual surgeon and the volume of the hospital are related to mortality\cite{4-8}. For example, one study showed that the 30-d mortality rate in centers performing > 21 resections per year was lower compared to centers performing ≤ 10 gastrectomies per year (4.4% vs 6.7%, $P = 0.047$)\cite{9}. Interestingly, more than half of the respondents from Asia perform > 60 gastrectomies.

The preferred method for gastric cancer surgery for most surgeons is an open gastrectomy. Only for early gastric cancer requiring a distal gastrectomy, the majority of surgeons preferably use a minimally invasive method. This is supported by recent short term results of the KLaSS-01 trial, which found that the complication rate was significantly lower after laparoscopic distal gastrectomy compared to the open distal gastrectomy (13% vs 20%, $P = 0.001$)\cite{10}. Evidence for all other types of gastric cancer are from combined with neoadjuvant chemotherapy in the majority of cases (73%). Differences in surgical approach and lymphadenectomy were found across continents.

In our survey, 79% of respondents performed

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**Figure 3** Open vs minimally invasive gastrectomy for early and advanced cancer.

**Figure 4** Lymph node dissection for distal and total gastrectomy.

**Table 1** Percentage of D2-dissections for different tumor stages worldwide (%)

| Region          | Asia  | Europe | South America | North America |
|-----------------|-------|--------|---------------|--------------|
| Distal early    | 33.9  | 57.1   | 54.2          | 38.5         |
| Total early     | 27.7  | 66.7   | 50.0          | 30.8         |
| Distal advanced | 98.2  | 85.7   | 91.7          | 92.3         |
| Total advanced  | 95.5  | 93.0   | 95.8          | 92.3         |

**Table 2** Percentage of different neo-adjuvant therapy regimens worldwide (%)

| Region         | Asia  | Europe | South America | North America |
|----------------|-------|--------|---------------|--------------|
| Chemotherapy   | 69.6  | 76.8   | 68.2          | 84.6         |
| Radiotherapy   | 0.0   | 1.8    | 0.0           | 0.0          |
| Chemoradiation | 12.5  | 12.5   | 9.1           | 7.7          |
| None           | 17.9  | 8.9    | 22.7          | 7.7          |
small randomized trials and retrospective studies only, but suggests minimally invasive gastrectomy to be safe.\textsuperscript{11-15} The absence of long-term results might explain the lack of generalized usage of these minimally invasive techniques. Randomized studies in both Asian and Western populations are awaited before worldwide implementation can take place.

In South Korea, the KLASS-02 trial and KLASS-03 (NCT01584336) trial are investigating the use of laparoscopy for distal gastrectomy for advanced cancer and total gastrectomy for early cancer respectively.\textsuperscript{16} Recently, in Europe two randomized controlled trials (LOGICA-trial and STOMACH-trial) started comparing open with laparoscopic gastrectomy.\textsuperscript{17,18} Regarding the current developments in minimally invasive surgery, one can expect the use of these techniques to increase. Since minimally invasive gastrectomy is associated with a considerable learning curve, expert training and proctoring is essential to provide for a safe implementation of this technique.\textsuperscript{19,20}

This survey showed that the majority of surgeons do not construct a jejunal pouch after total gastrectomy. Although literature remains scarce, studies have shown possible benefits of jejunal pouch reconstruction. Two studies demonstrated an improved quality of life in patients with a jejunal pouch, measured with the Gastro-Intestinal Quality of Life Index. In addition, they found no increase in postoperative morbidity after jejunal pouch reconstruction.\textsuperscript{21-24} Several trials on the use of a jejunal pouch are currently running to further investigate this possibly beneficial technique. This study demonstrated the preferable technique for constructing the anastomosis was by means of a mechanical stapler. Studies support these results, since a mechanical anastomosis is constructed significantly quicker compared to the hand-sewn method (11.4 min vs 38.7 min, $P < 0.001$) with a comparable complication rate.\textsuperscript{25,26}

For early cancers the majority of surgeons perform a D1+ dissection worldwide. For advanced tumors, the majority of surgeons perform a D2 dissection. The minority of surgeons performs resection of the spleen. These findings are in compliance with literature. In Western countries, pancreas and spleen preserving D2 dissection has been the preferred resection technique since two large trials from the Netherlands and the United Kingdom.\textsuperscript{27,28} These trials demonstrated D2 dissection with preserving the pancreas and spleen along with the associated lymph nodes (stations 10 and 11) to results in similar morbidity and mortality as D1 dissection and with better long-term results.\textsuperscript{27,29-31} In addition, it was argued that the reason for resecting station 10 and 11 is questionable since metastasis in these lymph nodes confers a poor prognosis (11 year survival: positive station 10 = 8%; negative station 10 = 27%; positive station 11 = 11%; negative station 11 = 35%).\textsuperscript{32} In Asia, surgeons perform a more tailored lymph node dissection. This survey showed that for early gastric cancer, a D2-dissection is performed less frequently in Asia compared to Europe and South America. On the other hand, the Japanese Gastric Cancer Guidelines still advise considering complete clearance of lymph node stations 10 by splenectomy for potentially curable T2-T4 tumors invading the greater curvature of the upper stomach.\textsuperscript{3} Some surgeons in Asia also perform a D3 dissection, since a Taiwanese trial showed an improved survival compared to D1 dissection.\textsuperscript{33} However, a D3 dissection did not improve survival compared to a D2 dissection in a Japanese trial.\textsuperscript{34} Evidence for a D1+ dissection is scarce. Only one small randomized trial demonstrated that D1+ dissection could be a safe alternative to D2 dissection for locally advanced non-junctional tumors. Therefore it seems justifiable that the majority of surgeons perform a D1+ dissection for early gastric cancer.\textsuperscript{35} More studies are needed to clarify the role of D1+ dissection for all types of gastric tumors.

The survey reveals that resection of the greater omentum was preferred by 88.5% of the participating surgeons. The value of resection of the greater omentum is currently debated. Advocates of its resection underline the importance of dissection of possible tumor deposits, whereas opponents argue that it is a time consuming procedure associated with additional morbidity. Literature on this topic is scarce and international guidelines vary. A retrospective cohort study in an Asian patient population demonstrated that the 3- and 5-year survival rates were not significantly different between gastrectomy with and without resection of the greater omentum for advanced gastric cancer.\textsuperscript{36}

Lastly, the majority of surgeons (84.4%) report the application of neoadjuvant chemotherapy or chemoradiation prior to gastrectomy. Literature presents various possible treatment strategies before and after surgery for gastric cancer. In Western countries, perioperative chemotherapy has shown the most beneficial, with an increase in survival around 13%.\textsuperscript{37} Interestingly, the percentage of surgeons in Europe using neo-adjuvant therapy in this study (2013-2014) is higher compared to the period of 2011-2012, where many patients did not receive neoadjuvant therapy.\textsuperscript{38} In Asia, adjuvant chemotherapy alone demonstrated better results compared to surgery alone with an increase in 3-year survival of 5%-10%.\textsuperscript{39-41} Adjuvant radiation in addition to perioperative chemotherapy can possibly increase survival and is currently investigated in Western countries.\textsuperscript{42,43} In Asia however, adjuvant radiation in addition to adjuvant chemotherapy did not increase 7-year overall survival (75% vs 73%, $P = 0.484$).\textsuperscript{44}

A limitation of this study is that it only evaluates expert opinions rather than objective measurements, which should be taken into account before generalizing these findings. However, its international design provides a unique insight in the current practice of gastric cancer surgeons. Its discussion in the light

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of contemporary literature can be used for further improvement of gastric cancer surgery worldwide. Lastly, these results can be used for future evaluation of worldwide gastric cancer surgery.

In conclusion, this study is unique in its international design revealing the expert opinion on gastric cancer surgery. Minimally invasive gastrectomy is still not widely adapted and variations between continents are present. Minimally invasive gastrectomy is most popular in Asia to treat patients with early gastric cancer. Neoadjuvant chemotherapy is used by the majority of surgeons worldwide. A D1+ lymphadenectomy is preferred for early gastric cancer and a D2 lymphadenectomy is preferred for advanced gastric cancer.

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