Perforated gastric and duodenal ulcers: treatment options

Abstract
This paper focuses on a prospective nonrandomized review of patients undergoing surgery for perforated gastric and duodenal ulcers.

Patients and methods: A total of 198 patients with perforated gastric and duodenal ulcers were enrolled in the study between 2011 and 2016. The mean age of patients was 42 years. The disease was more common in men (87.3%) than in women (12.6%).

The incidence of duodenal ulcer perforation was 86.3%. Anti-helicobacter therapy was administered to 33.8% of patients before perforation. In 5.6% of cases, recurrent ulcer perforation was established. The majority of patients (78.8%) were admitted within the first 12 hours, while 7.5% - 24 hours after perforation. The APACHE II scoring system was used to measure shock in 5.5% of cases. A score up to 6 was determined in 37.8% of patients, up to 12 - in 47.9%, higher scores (more than 12) were measured in the remaining patients. Physical examination and diagnosis included clinical methods, abdominal X-ray, gastroduodenoscopy, ultrasound, laparoscopy, and the Boey risk scores.

Results: Diagnostic laparoscopy was performed in 79.3% of patients, the diagnosis of concealed perforation was confirmed by gastroduodenoscopy. Video-assisted laparoscopy was performed in 59% of patients, 14.7% - underwent combined minimally invasive surgery (abdominal laparoscopic examination and surgical intervention in the stomach or duodenum using a minimally invasive technique) and 26.3% - via laparotomy. Laparoscopic surgery was possible on the basis of gastroduodenoscopy, laparoscopy (perforation diameter, a correlation between the size of ulcer and perforation diameter, perulcerous infiltration sizes). Postoperative complications were recorded in 8.1% of cases, the mortality rate was 4.5%.

Keywords: perforated gastric and duodenal ulcer, diagnostics, surgical treatment

Introduction
Perforated gastric and duodenal ulcers (PGDU) continue to be one of surgical problems accompanied by a 30% mortality rate and complications up to 50% including an acute abdominal syndrome, limited or generalized peritonitis and a high risk of sepsis.1 of the complications encountered, perforation ranks second in frequency after bleeding (the ratio is 1:6) and is the most common indication for emergency surgery for peptic ulcer.2-4 In 2006, more than 150,000 patients with peptic ulcer (PU) complications were hospitalized in the USA alone. The total number of complications due to perforation (n=14500–9%) was as much as 7 times less than bleeding. Peptic ulcer perforations caused death in 37% of cases5 and more than one in every admission related to peptic ulcer complicated by perforation was fatal. In ulcer perforation, the mortality rate was about 5 times higher than that in bleeding from the site of perforation. It was also the most significant death factor in US hospitals between 1993 and 2006 (OR: 12.1 (95% CI: 9,8-149)).2

Over the past 30–40 years, the European mortality rate associated with PGDU has been stable despite some progress made in perioperative management, visualization and surgical treatment techniques.5-7

An imbalance between protective and ulcer-promoting factors in the pathogenesis of PU is evident. However, the causes of perforation in some patients need to be elucidated. Ulcerogenesis is associated with infections (H. pylori), disruptions in the mucosal protective barriers (the administration of certain drugs), and the excessive secretion of acid, although appropriate risk assessment and the impact of each factor are not sufficiently studied.8 Only about one third of patients with PGDU have a history of PU. In some patients, small perforations (<5 mm in diameter) without large mucosal defects occur. This fact may suggest that the size of an ulcer is not related to perforation risk development, although in a number of patients with large defects (a few cm in diameter) perforations may occur. The latest development has been the recognition of the role of Helicobacter pylori in the pathogenesis of perforation.9,10

A total of 198 patients with PGDU were treated at the Ufa Emergency hospital between 2011 and 2016. The mean age of patients was 42 years. The disease was more common among males - 173 (87.3%) than females - 25 (12.6%). Perforated duodenal ulcers were detected in 171 patients (86.3%), gastric localization was found in 27 patients (13.7%).

Perforated duodenal ulcers occurred most commonly among young patients, whereas gastric ulcers - in the aged. Of the total number, 8
patients (4%) were given nonsteroid anti-inflammatory drugs before perforation. Of interest, before admission related to PGDU anti-ulcer therapy was administered to 184 patients (92.9%). Among them, 67 patients (33.8%) received anti-helicobacter therapy. Out of them, only in one fifth of patients the treatment outcomes were assessed by fibrogastroduodenoscopic diagnostic testing for the presence of H. pylori. Eleven patients (5.6%) underwent surgery for recurrent gastric ulcer perforations (4) and duodenal ulcer perforations (7). Perforation combined with other complications was encountered in 6 cases (3%) including gastroduodenal hemorrhage in 1 patient (0.5%), stenosis of the pylorus in 1 case (0.5%), ulcer penetrated into the head of the pancreas and small omentum in 4 cases (2%). The following distribution of admissions was established: 71.2% of patients were admitted within 6 hours after disease, 15 (7.6%) - from six to twelve hours, 27 (13.6%) - from 12 to 24 hours, 12 (6%) - from 24 to 48 hours, and 3 patients (1.5%) had very delayed presentation (over 48 h). Shock on admission was recorded in 5.5% patients.

The APACHE II scoring system was used to assess patients’ illness severity: a score of up to 6 was in 75 patients (37.8%), up to 12 - in 95 (47.9%), from 12 to 18 - in 22 (11.1%), and over 18 - in 6 patients (2.8%).

In PGDU diagnostics, along with abdominal radiography fibrogastroduodenoscopy was used. When necessary, re-radiography following endoscopy, ultrasound of the abdomen and diagnostic laparoscopy could be applied. Laparoscopy was performed in 79.3% of patients.

We faced certain diagnostic difficulties in patients with concealed peptic ulcer (n=11 patients). In these cases the diagnosis was established ultrasonographically and gastroduodenoscopy. In this patient group, the time of final updated diagnostics did not exceed 3 hours. In the remaining patients the diagnosis was verified within the first two hours after admission.

Endoscopic diagnostics - gastroduodenoscopy is essential in determining the location and diameter of the perforation. The following criteria are the most applicable to choose the perforation suture:

1. The size of ulcer.
2. The perforation diameter.
3. Periulcerous infiltration sizes.
4. The peritonitis extent.

We determine the location of the ulcer and its size, perforation size using gastroduodenoscopy. Occasionally there may be difficulty in determining the periulcerous infiltration size. Although laparoscopy is believed to be a more preferable technique, but with marked tissue infiltration it is hardly helpful for suturing large perforations. A laparotomic approach was applied with perforations over 10 mm, and 3 patients (1.5%) had very delayed presentation (over 48 h). Shock on admission was recorded in 5.5% patients.

The indications for laparatomy were primarily established in 8 patients with generalized peritonitis, a critical state according to the APACHE II scores (>18 and over), delayed presentation of patients (>24 h from onset).

According to the literature data, the frequency of conversion to laparotomy is 28.5%, average is about 12.4%. Conversion is recommended in the presence of technical difficulties related to suture

Discussion

Despite the availability of long-term experience in treating patients with PGDU, some issues remain disputable. It is necessary to reveal correlating risk factors for improving potential treatment outcomes. Thus, the most widely used prediction model is the Boey clinical score based on the presence of serious diseases, preoperative shock and long standing perforation (>24 h).12–14 However, its positive prognostic value in 94% of cases was not confirmed by subsequent studies.12–14

Other prognostic criteria specifically for PGDU were proposed.11 Nevertheless, none of them was confirmed in external cohorts. This prevents their generalization. In addition, some common surgical systems and scores were used.

The authors’ technique based on the correlation between the size of the ulcer and perforation diameter simplifies the treatment option and reduces the number of complications. So, complications in the patient group operated on laparatomically were higher (OR: 0.240 (95%), CI: 0.063-0.656) compared with laparoscopic interventions. With patient randomization, if the coefficient was less than 2 and without it (44 patients with the coefficient and 73 without it) complications were significantly lower (1 and 3 complications, respectively) (OR:0.543 (95%) CI 0.055-5.385). Similar results were obtained in patients with an open procedure (OR:0.210 (95%) CI 0.042-1.042): in the presence of the coefficient, there were 2 complications in 38 patients, whereas without it - 9 complications in 43 patients.

All 198 patients with PGDU underwent surgery predominantly within the first two hours after admission. Patients with marked intoxication, unstable hemodynamics, signs of peritonitis, infectious and toxic shock (n=22 patients) needed short-term preoperative management. The indications for management of these patients included central venous pressure up to 8-12 mm Hg, mean arterial pressure > 65 mm Hg, Svo₂>70%, diuresis > 0.5 ml/(kg*h).

Patients with PGDU were treated using three surgical interventions: a video-assisted laparoscopic approach (n=117 (59%)); combined minimally invasive approaches - abdominal laparoscopic examination and interventions in the stomach or duodenum using a mini-laparotomic approach (n=29 (14.7%)) and surgery using a routine laparotomic approach (n=52 (26.3%)).

The indications for laparotomy were primarily established in 8 patients with generalized peritonitis, a critical state according to the APACHE II scores (>18 and over), delayed presentation of patients (>24 h from onset).
of the perforation site, when the ulcer location is not demonstrable and there are large perforations (>10 mm).\textsuperscript{15,16} The use of omenoplasty may reduce the possibility of conversion and with perforations over 10 mm.

At surgery, concealed perforation was revealed in 11 patients (5.5%), localized peritonitis in 29 (4.6%), diffuse in 32 (16.1%) and diffuse peritonitis in 126 patients (63.6%). According to clinical criteria, the reactive phase of peritonitis was established in 47 patients (23.7%), toxic - in 142 (71.7%) and terminal - in 9 patients (4.5%). Laparoscopic approaches have a number of advantages to most patients with PGDU. On these bases they were performed in 146 patients (73.6%) including combined ones with ulcer suturing using a mini-laparotomic approach in 29 patients (14.6%).

Contraindications to laparoscopy in PGDU were considered to be the patient’s critical health state (the total APACHE scores >18), terminal peritonitis phase, diffuse peritonitis with multiple abdominal abscesses, the necessity in intubation and decompensation of the bowel, generalized peritonitis along with surgery for the abdominal cavity organs. In the authors’ view, additional objective assessment criteria for laparoscopy are the grade of intra-abdominal hypertension syndrome and the level of abdominal perfusion pressure (determined as a difference between average arterial and intra-abdominal pressure). Critical criteria are an increase in intra-abdominal pressure >20 mm Hg and abdominal perfusion pressure < 65mm Hg. The most safe is laparoscopy with the Boye risk score of 0-1.\textsuperscript{11}

At combined minimally invasive surgery, laparoscopic examination of the abdomen was initially performed, and then suture of the perforation site was done using a mini-laparotomic approach. This type of surgery was applied when there were technical difficulties with laparoscopic suture, the large size of perforation, marked infiltration changes around the perforation site, organ abnormalities, when it was impossible to suture the perforation and in the presence of combined complications (perforation + bleeding or + stenosis, etc.).

Grade I perforated ulcer (the perforation diameter < 2mm) was seen in 4 patients (2%), Grade II (diameter 2-5 mm) - in 96 patients (48.5%), Grade III (diameter 6-10 mm) - in 75 patients (37.8%) and Grade IV (diameter >10mm) - in 22 patients (11.1%).

In our clinic, suture of the perforation site was performed in 191 patients (96.5%), suture in conjunction with proximal selective vagotomy - in 5 (2.7%) and gastric resection - in 2 patients (1%). To suture, one-row vicril sutures (n=88-46%), two-row (n=71 – 38.7%), sometimes suture with two-row sutures with ulcer resection (n=18 – 9.4%) and two-row sutures with omenoplasty (n=11 – 5.7%) were used.

One-row sutures were used for small perforation sites (grades I-II) and in the absence of marked inflammatory changes around the ulcer. Two-row sutures were used for perforation sites of grades III-IV and in the presence of inflammatory changes around the ulcer. Two-row sutures with omenoplasty were used for perforation sites of grades III-IV with marked changes (inflammatory wall >10mm) as a rule applying laparotomic or mini-laparotomic approaches.

Suture with ulcer defect resection was used with pyloroduodenal localization due to the eruption of sutures, stenosis of the lumen of the pylorus or the duodenum, as well as when pyloplasty was necessary. Mandatory gastric resections were performed in one case when malignancy of the gastric localization was suspected. The second case was associated with the presence of a giant ulcer and impossibility to suture the perforation site.

Postoperative complications developed in 16 patients (8.1%) including failure of sutures in one case (0.5%), intra-abdominal abscesses - 6 (3.1%), suppuration of operation wounds in 9 cases (4.5%). After laparoscopic suture and examination, the suture failure was seen in 2 cases, after laparotomy - in 4 cases (OR:0.104 (95% CI 1.063-1.130). Suppuration of wounds occurred after an open procedure (8), in one case - after mini-laparotomy.

Postoperative lethality in the study group was 4.5% (9 patients). The causes for unfavorable outcomes were septic shock, uncured diffuse peritonitis, multi-organ insufficiency, thromboembolism of the lung artery. All those patients who died had undergone surgery using a classical laparotomic approach in the state of shock due to diffuse peritonitis with intra-abdominal abscesses. All of them were admitted after 24 hours after onset, and 3 patients after 48 hours.

Conclusion

Perforated gastric and duodenal ulcers remain a challenge in modern surgery due to high rates of complications and mortality. At the present time the issues of the pathogenesis of gastroduodenal ulcer perforation, mechanisms of regeneration as well as the risk of sepsis need to be better considered. In cases when the perforation is large (>20mm) or inflamed tissues are friable there is difficulty in achieving safe healing of the wound by a primary suture. If the first attempt of the defect closure fails, the next one may also result in failure. In such cases resection should be considered, taking into account that large gastric ulcers or recurrent perforations must serve as an important indicator of malignancy which occurs in about 30% of patients.\textsuperscript{16,17} In these circumstances, gastric resection, partial resection of the stomach with gastrounostomy or insertion of a T-like drainage should be conducted if the defect is located in the duodenum.\textsuperscript{16}

In recent years, to treat PGDU the endoscopic application of clips or stenting,\textsuperscript{19,20} the use of biodegradable material to close the defect in the ulcer site,\textsuperscript{21} mesenchymal stem cells to improve healing have been proposed.\textsuperscript{22-27}

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Conflicts of interest

The authors declare there is no conflict of interest.

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