Duodenal and jejunal Dieulafoy’s lesions: optimal management

Tonguç Utku Yılmaz¹
Ramazan Kozan²
¹Acibadem University Atakent Hospital, Organ Transplantation Unit, Istanbul, Turkey, ²Eren Hospital, General Surgery Unit, Istanbul, Turkey

Abstract: Dieulafoy’s lesions (DLs) are rare and cause gastrointestinal bleeding resulting from erosion of dilated submucosal vessels. The most common location for DL is the stomach, followed by duodenum. There is little information about duodenal and jejunal DLs. Challenges for diagnosis and treatment of Dieulafoy’s lesions include the rare nature of the disease, asymptomatic patients, bleeding symptoms often requiring rapid diagnosis and treatment in symptomatic patients, variability in the diagnosis and treatment methods resulting from different lesion locations, and the risk of re-bleeding. For these reasons, there is no universal consensus about the diagnosis and treatment approach. There are few published case reports and case series recently published. Most duodenal DLs are not evaluated separately in the studies, which makes it difficult to determine the optimal model. In this study, we summarize the general aspects and recent approaches used to treat duodenal DL.

Keywords: Dieulafoy’s lesion, gastrointestinal bleeding, duodenum, endoscopy

Introduction

Dieulafoy’s lesions (DLs) are most often located in the stomach; also, they have been detected in the duodenum. While gastric DLs are well known, there are few reports about duodenal and small intestinal lesions. Before endoscopy was used, the diagnosis and prognosis for patients with these lesions was poor, with mortality rates ranging from 23% to 79%. The probability of forward-viewing endoscopy failure in the diagnosis of duodenal DL, the high-risk surgery for periampullary DL, and re-bleeding risks made the treatment and the diagnosis difficult for duodenal lesions. Similarly, localization problems in jejunal DLs prompted clinicians to search for new diagnostic and treatment modalities.

Duodenal and jejunal DLs constitute 15% and 1% of all DLs, respectively. Duodenal and jejunal DLs are responsible for 3.5% of all gastrointestinal (GI) bleedings. Unrecognized lesions and the rare nature of their occurrence make these lesions difficult to diagnose, and it is also difficult to develop a standardized treatment for patients with DLs. Duodenal DLs are rare and they are mostly presented as case reports. In this study, we report the most recent information about the diagnosis and treatment modalities for patients with duodenal and jejunal DLs from the limited number of studies.

Pathology and etiology

DLs are different from typical peptic ulcers because the mucosal defect in DLs is not surrounded by inflammatory cell infiltration, and the exposed artery has a large...
Clinical presentation and diagnosis

The clinical presentation of DLs is painless with massive periodic recurrent intermittent hematemesis associated with melena, hematochezia, and hypotension. Concomitant peptic ulcer or gastritis might present with previous dyspeptic symptoms. While duodenal DL patients mostly present with hematemesis, jejunal DL patients mostly present with melena. Hemodynamic stability is important in making a diagnosis and deciding upon treatment modalities. The clinical DL findings are similar to GI bleeding and include hematochezia, melena, hematemesis, tachycardia, hypotension, and syncope. Intermittent and massive GI bleeding is included in the classical DL presentation. Patients presented with melena (44%), hematemesis (30%), melena and hematemesis (18%), and hematochezia and iron-deficiency anemia (6%). The mean blood hemoglobin levels in patients presenting with DL ranged from 4 to 14 g/dL. The clinical presentations of patients with DL depends on the duration of bleeding, general condition of the patient, localization of bleeding, and diameter of the bleeding vessel. While patients with duodenal lesions generally present with symptoms of upper GI bleeding, patients with jejunal DL present with symptoms of lower GI bleeding (Table 1).
Jejunal DLs are very rare and constitute only 3.5% of all small intestinal bleeds. There were <50 patients with jejunal DLs in the literature. Jejunal DLs are not in the range of standard endoscopies, except the proximal ones. Thus, push, single-, or double-balloon enteroscopy can be used for both diagnosis and treatment. Alternatively, capsule endoscopy, angiography, and red cell scanning can be used to pinpoint the location of the jejunal DL. Angiography and red cell scanning can be used to determine the location of the bleed. There is no specific radiological view of the lesion. The angiographic findings include extravasation of contrast from an eroded artery that may appear normal. However, existence of a tortuous and ectatic artery in angiography can provide important information for the diagnosis of DL. Although the patients’ hemodynamic status is important for deciding on diagnostic modalities, most of the patients were evaluated first by colonoscopy or endoscopy for the evaluation of GI bleeding. Technetium scintigraphy was used in 2 studies but it was unable to show any bleeding site. Angiography can show bleeding points that originate from the mesenteric arteries. If standard angiography fails, formal heparin therapy with an angiogram can be used to show the exact bleeding point. The selective placement of an angiographic catheter into the affected artery may also have resuscitative and possible therapeutic benefits. Intra-arterial vasopressin installation or embolization may aid in slowing any hemorrhaging, thereby allowing more time for further resuscitative efforts. Patients who undergo embolization are at risk of ischemia of the non-involved jejunal segments unless the catheter has been placed distal to the subsegmental branches. Angiographic embolization is the choice of treatment in suitable patients. Angiography failed to show the bleeding point in 2 of 4 jejunal DL patients, and embolization was also unsuccessful in these patients. The most likely reason was the relative mobility of the jejunum and the thinness of its wall, both of which made it difficult to obtain the desired compression of the vessel by angiographic embolization. Patients who underwent successful embolization for the treatment of DL were also shown to have another DL in another organ. Capsule endoscopy is a frequently used method to evaluate bleeding in the small intestine. It is a method of choice for small intestine imaging after colonoscopy and endoscopy for mid-GI bleeding in hemodynamically stable patients. Capsule endoscopy has pinpointed the origin of the bleeding in all patients with jejunal DL. Localization of the bleeding point guides the clinician in the best way to approach the lesion. The decision as to whether the initial enteroscopy was performed from an oral or anal approach was based

<table>
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<th>Table 1</th>
<th>Age, sex, presenting symptoms, and average number of endoscopies needed prior to diagnosis for both duodenal and jejunal DLs</th>
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<td><strong>Demographic data</strong></td>
<td><strong>Duodenal DL</strong></td>
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<td>Mean age (years), (range)</td>
<td>67.08 (21–86)</td>
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<td>Sex predominance</td>
<td>No</td>
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<td>Most common presentation</td>
<td>Hematemesis</td>
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<td>Mean average number of endoscopies for diagnosis (range)</td>
<td>1.8 (1–8)</td>
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<td>Rate of re-bleeding after endoscopic treatment (%)</td>
<td>0–10</td>
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Abbreviation: DL, Dieulafoy’s lesion.

In the past, the diagnosis was rarely made before surgery or postmortem examination. The diagnosis could only be made by histological/pathological examination. However, with increased surgical experience, the diagnostic criteria are now dependent on endoscopic findings. Endoscopic criteria for DLs include several parameters: 1) active arterial spurting or micropulsative streaming from a tiny mucosal defect or through the normal surrounding mucosa; 2) visualization of a protruding vessel with or without active bleeding within a tiny mucosal defect or through the normal surrounding mucosa; and/or 3) fresh, densely adherent clot(s) with a narrow point of attachment to a tiny mucosal defect or to normal appearing mucosa. Currently, DL is diagnosed endoscopically rather than histologically. Upper GI endoscopy can easily reach the duodenum and help to diagnose and treat DL. However, the diagnosis may be difficult to make because of the small nature of the lesion, the normal appearance of the surrounding mucosa, and the intermittent nature of the hemorrhages. Therefore, multiple endoscopies are often necessary to make the diagnosis. In addition to the nature of the lesion, massive hemorrhage in the limited lumen of the duodenum can be overlooked because of other concomitant lesions such as an ulcer, and an inexperienced endoscopist who looks for an inflamed lesion can overlook the DL because of clots, duodenal angulations, or periampullary diverticulum. Thus, the duodenal DL cannot always be controlled on the first attempt. Lopez-Arce et al showed that patients underwent at least 3 attempts to control the duodenal DL bleedings. To overcome this problem, early endoscopic evaluation and side-viewing endoscopy is needed. Endoscopies performed within the first 12 hours have a high success rate for diagnosing DLs because of their capability to pinpoint the bleed location. Duodenal DLs located in the periampullary area and in the second portion of the duodenum cannot be seen or treated by forward-viewing endoscopies. Barium studies were unsuccessful in duodenal DL diagnosis.
primarily on clinical presentation and the result of capsule endoscopy. If capsule endoscopy reveals a possible bleeding source within the proximal two-thirds of the small bowel, an oral route enteroscopy is performed first because it can reach the bleeding point. If no bleeding is detected along the oral route, a submucosal tattoo is placed to mark the deepest insertion point and the anal route is then used. Tattooing can also be used for marking the lesion(s) during surgery when endoscopic treatments fail. Tattooing the bleeding site with India ink maybe helpful for future identification in the event of recurrent bleeding. Single- or double-balloon enteroscopy is used to treat mid-GI bleeds. Chronic intermittent bleeding may be encountered when treating jejunal DL, which results in multiple balloon-assisted enteroscope procedures before diagnosis; it was reported that at least 2 or more enteroscope procedures were needed before diagnosis.19,24 Although Dulic-Lakovic et al reported that 1.5 endoscopies were required per patient to diagnose jejunal DL, Lipka et al reported that all jejunal DL patients were diagnosed after the first attempt.8,19 Single- or double-balloon enteroscopy is mostly preferred method for treatment of jejunal DLs in recent studies. If the patient is hemodynamically unstable or if technical experience with endoscopic treatment is limited, surgical evaluation is needed. Intraoperative enteroscopy from an enterotomy can help to focus on the area containing the bleed.13,21 Lipka et al also showed that DLs were localized in the proximal jejunum.8 Dulic-Lakovic et al showed that the mean distance of the jejunal DL from the pylorus was 132±115 cm.19 For both duodenal and jejunal DLs, DLs in other accompanying organs should be considered.22

**Treatment**

DLs are a cause of GI bleeding, and treatment depends on the presentation, lesion site, and available surgical and endoscopic expertise. Bleeding from DLs may be life threatening and they were treated surgically before 1990. Since then, endoscopic treatment has replaced surgery as the treatment of choice. Several endoscopic treatment methods, including injection, a heater probe, bipolar electrocoagulation, and laser therapy have been used. Since then, mechanical devices such as hemoclips and rubber band ligation have been presented as new choices for the treatment of GI bleedings.

Injection therapies mainly aim to stop bleeding from vessels by injection of several agents such as vasoconstrictors (epinephrine) or sclerosants (ethanol, aethoxysklerol, or polidocanol).17 One DL patient who was treated with N-butyl 2-cyanoacrylate has been described in the literature.25 Epinephrine can be injected (dilution of 1:10,000) into 4 quadrants, with an initial dose of 2.5 mL that can be repeated. This is an inexpensive, simple, and relatively safe procedure. There is the risk, however, of re-bleeding. In 1 case, epinephrine was used as the sole therapy in duodenal DL without re-bleeding. However, it has been shown that epinephrine plus a heat probe is the most preferred treatment method in duodenal DL.6 Initial injection with epinephrine, followed by other treatment methods, seems like an optimal treatment for acute bleeding because it provides the advantages of temporary hemostasis without risk of tissue injury and perforation. Achieving enhanced visualization of the bleeding site should guide the second definitive treatment method (such as a combination of mechanical approaches) and reduce the risk of complications.

Thermal endoscopic treatments stop the bleeding using heat either by contact or non-contact. Bipolar and heater probes are examples of contact thermocoagulation, and argon is an example of a non-contact method. Contact thermocoagulators carry a risk of transmural injury in thin walled organs such as the duodenum. The non-contact methods have less tissue depth injury, and they are safer than the contact methods. However, they are only effective for superficial coagulation. There is no reported case of perforation after duodenal DL treatment with a heater probe. Schmulewitz et al presented 3 cases in which duodenal DLs were treated only with heater probes.6

Mechanical hemostasis is performed using hemoclips and band ligation. Hemoclips have been suggested to be more successful than thermal ablation in achieving hemostasis for bleeding DLs. The success rate using hemoclips is generally high when the surrounding tissue is soft. Thus, hemoclips are advantageous for hemostasis in DLs because the size of the lesion is small, and hemostasis can be obtained by concurrently clipping the normal surrounding mucosa. It is difficult, however, to apply hemoclips in difficult locations such as the duodenum; acute duodenal angulation makes hemoclip application maneuvers impossible. Side-view endoscopes, which require surgical expertise, may be needed for hemoclip application in difficult locations.12 Hemoclips that were previously placed incorrectly can hinder accurate positioning of subsequent hemoclips. Thus, it is important to apply the hemoclips accurately to the DL the first time they are used. Epinephrine injection(s) before hemoclip application might be helpful in facilitating this procedure.

Band ligation is easier than hemoclip application because of its accessibility to difficult locations. It is safer than thermal techniques in terms of minimizing perforations. There is a risk, however, of re-bleeding ulcer formation around the ligated mucosa. Another problem is the required time interval...
for preparing the overtube and other pieces of equipment for band ligation. Chung et al compared the efficacy of hemostatic methods in patients with DL, and they reported that mechanical methods for hemostasis such as band ligation and hemoclipping were superior to injection methods for controlling bleeding and preventing recurrent bleeding. In duodenal DLs, however, mechanical techniques are difficult and injection plus thermal ablation achieves higher success rates with less re-bleeding.

Based on these advantages of endoscopic treatment, nearly all patients with DLs can be treated with endoscopic methods; primary hemostasis may occur in more than 90% of patients. Re-bleeding has been found and has not been associated with age, sex, medical history, or location. Duodenal localization does not appear to present a risk factor for re-bleeding. Monotherapy with adrenaline and arterial spurting are risk factors for re-bleeding. For this reason, combined therapy is advised for duodenal DLs. Park et al showed that re-bleeding was not associated with DL location or treatment method. The presence of infection and kidney disease were related to re-bleeding.

If endoscopic equipment is lacking or endoscopic treatment of duodenal DLs has failed, exploratory surgery must be performed. The rate of surgical necessity for DLs has decreased from 100% to 3% with increasing surgical experience. If the localization of the duodenal DL is known, it is preferable to longitudinally open that part of the duodenum. If the location is not known because there is no predominant localization of the duodenal DL, it would be better to longitudinally open the anterior wall of duodenal bulb. This would help the surgeon evaluate the papilla, any possible ulcers, and the distal part of the duodenum. The bleeding point can be overlaid with a 3-point U stitch technique. Care must be taken to avoid incorporating the common bile duct into a stitch. The course of the common bile duct can be identified by inserting a probe through the ampulla. After bleeding is controlled, the duodenotomy should be closed transversely to avoid narrowing. Unless pylorotomy is performed, there is no need for vagotomy. If the source of bleeding cannot be defined, emergent pancreaticoduodenectomy should be performed.

Patients with DL usually present with GI bleeding. Therefore, the patients should first be treated using a general GI bleeding approach. In these cases, necessary fluid replacement, blood transfusions, and blood products, and close follow-up monitoring should be provided. The main factor in deciding which treatment modality to use is patient’s hemodynamic status. If the patient is hemodynamically unstable, exploratory surgery must be performed. However, without localization of the bleeding point, exact resection of the jejunal DL cannot be performed on the first attempt. Because DLs do not have any fibrosis, inflammation, or surrounding polyps, it is nearly impossible to feel the DL. For this reason, intraoperative endoscopy (oral, anal, or via enterotomy) is needed (Figure 1). Historically, intraoperative methylene blue injection via the selectively positioned catheter can be used to identify the affected jejunal segment. Segmental resection and anastomosis is the preferred surgical choice to treat jejunal DL (Figure 2). Simple overlaying of the lesion is not recommended because it is associated with a greater risk of recurrent bleeding. There was no re-bleeding in surgically treated patients. Jejunal DL bleeding is generally obscure, and early treatment is needed. The primary modality of endoscopic treatment is electrocautery and multipolar electrocoagulation. Epinephrine injection is used as an adjuvant therapy to initially slow down the bleeding. If coagulation is unsuccessful, clip placement is used for hemostasis. Argon plasma coagulation is the first choice to treat patients with
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<th>Localization</th>
<th>Age, years</th>
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<th>Number of endoscopies for treatment</th>
<th>Treatment</th>
<th>Re-bleeding</th>
<th>Follow-up</th>
<th>Failed treatment modalities</th>
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<td>36</td>
<td>M</td>
<td>Melena, hematochezia</td>
<td>Intraoperative enterostomy</td>
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<td>After 15 days</td>
<td>NM</td>
<td>Endoscopy, capsule endoscopy, angiography, intraoperative enteroscopy</td>
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<tr>
<td>J</td>
<td>13</td>
<td>M</td>
<td>Hypovolemia</td>
<td>1</td>
<td>Laparoscopic bowel resection after capsule endoscopy</td>
<td>After 15 days</td>
<td>NM</td>
<td>Endoscopy, scintigraphy</td>
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<tr>
<td>J</td>
<td>35–82</td>
<td>M/F:4/6</td>
<td>Obscure bleeding</td>
<td>1 in six patients 2 in three patients 3 in one patient</td>
<td>Argon in three patients Clips in three patients Injection in one patient Injection + clip in one patient Injection + argon in one patient Bowel resection in one patient</td>
<td>1 patient after 13 days 1 patient after 49 and 67 days</td>
<td>10–17 months</td>
<td>I patient with argon and injection I patient with argon and secondly with clips</td>
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<td>J/D: 7/1</td>
<td>60–86</td>
<td>M/F:4/4</td>
<td>Melena</td>
<td>1 in three patients 2 in four patients 3 in one patient</td>
<td>Bipolar in two patients Bipolar+argon+clip in one patient Bipolar+argon+clip in one patient Bipolar+argon+clip in two patients Argon in one patient Epinephrine + bipolar in one patient</td>
<td>After 5 days</td>
<td>NM</td>
<td>Endoscopy, angiography, laparotomy, intraoperative enteroscopy Endoscopy, capsule endoscopy, angiography Endoscopy, capsule endoscopy, enteroscopy, subtotal colectomy</td>
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<td>J/D: 7/1</td>
<td>67–76</td>
<td>F</td>
<td>Melena</td>
<td>Multiple</td>
<td>Epinephrine and clip in one patient Bowel resection in one patient Argon in one patient Laparoscopic bowel resection after capsule endoscopy in one patient</td>
<td>After 5 days</td>
<td>NM</td>
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<td>Holleran et al</td>
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### Table 2 (Continued)

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<thead>
<tr>
<th>Localization</th>
<th>Age, years</th>
<th>Sex</th>
<th>Symptoms</th>
<th>Number of endoscopies for treatment</th>
<th>Treatment</th>
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<th>Follow-up</th>
<th>Failed treatment modalities</th>
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<tr>
<td>J</td>
<td>25</td>
<td>M</td>
<td>Hematochezia</td>
<td>1</td>
<td>Intraoperative enteroscopy</td>
<td>7 and 14 days</td>
<td>Angiographic embolization 2 times</td>
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<tr>
<td>J</td>
<td>21</td>
<td>F</td>
<td>Hematochezia</td>
<td>Intraoperative enteroscopy</td>
<td>Bowel resection</td>
<td>+</td>
<td>24 months</td>
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<tr>
<td>J</td>
<td>54</td>
<td>M</td>
<td>Hematochezia</td>
<td>Intraoperative enteroscopy</td>
<td>Segementer resection</td>
<td>+</td>
<td>24 months</td>
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<tr>
<td>J</td>
<td>63</td>
<td>F</td>
<td>Melena</td>
<td>2</td>
<td>Oral push endoscopy</td>
<td>2 months</td>
<td>Angiography and scintigraphy</td>
<td>Eddi et al(^1^5)</td>
<td></td>
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<tr>
<td>J</td>
<td>20</td>
<td>F</td>
<td>Hematochezia</td>
<td>Intraoperative enteroscopy</td>
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<td>15 days</td>
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<tr>
<td>J</td>
<td>36</td>
<td>M</td>
<td>Hipovolemia</td>
<td>1</td>
<td>Intraoperative enteroscopy and bowel resection</td>
<td>2 months</td>
<td>NM</td>
<td>Goins et al(^2^1)</td>
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<tr>
<td>D:7</td>
<td>44–84</td>
<td>–</td>
<td>–</td>
<td>1–4</td>
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<td>6 months</td>
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<tr>
<td>D</td>
<td>74</td>
<td>M</td>
<td>Melena</td>
<td>8</td>
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<td>NM</td>
<td>6 months</td>
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<tr>
<td>D</td>
<td>25</td>
<td>F</td>
<td>Melena</td>
<td>1</td>
<td>Epinephrine</td>
<td>NM</td>
<td>7 months</td>
<td>Ibrarullah et al(^1^1)</td>
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<tr>
<td>D:2</td>
<td>76, 77</td>
<td>M/F</td>
<td>Hematemesis, melena, AMI</td>
<td>NM</td>
<td>Epinephrine and coagulation</td>
<td>72 hours one patient</td>
<td>NM</td>
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<tr>
<td>D</td>
<td>64–85</td>
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<td>One patient rebleed</td>
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Note: "+" means positive re-bleeding but no detailed information was given.

Abbreviations: AMI, acute myocardial infarction; D, Duodenum; F, Female; J, Jejunum; M, Male; NM, not mentioned.
jejunal DLs. There were 4 reported cases of re-bleeding in jejunal DL patients after endoscopic treatment. The rate of re-bleeding after endoscopic treatment of jejunal DL is given as 12.5% and 20% in previous studies. The re-bleeding times were 12, 13, and 49 days and 12 weeks after the first treatment. The primary hemostasis treatments were argon and clips. There is not enough data to permit comparison of the treatment modalities. Studies have shown that mechanical endoscopic methods such as hemoclip and band ligation are more effective than injection and thermal therapy for general DL. Additionally, it has been shown that using a combination of 2 endoscopic techniques for hemostasis is superior to using a single method for treating gastroduodenal ulcer disease. Therefore, optimal endoscopic treatment of jejunal DL would be to combine thermal therapy and clip application. Mortality after jejunal DL bleeding depends on existing comorbid diseases such as cardiovascular diseases or cirrhosis. For this reason, early diagnosis and treatment is important. The mortality rates described in previous studies, in which endoscopic treatment was not popular, were between 23% and 35%. With increased surgical experience in endoscopic treatment and awareness of DL, early diagnosis has decreased mortality rates by up to 8%. While the mortality rates in previous studies were associated with massive bleeding, recent mortality rates were mostly associated with comorbid diseases such as heart and/or respiratory failure, sepsis, and/or cirrhosis.

When re-bleeding occurs, diagnostic and treatment approaches were the same as the previous treatment approaches. As there is a risk of re-bleeding in DL patients even 12 weeks after the initial treatment, the clinician should follow-up with the patient for at least 6 months. A longer follow-up does not add any benefit for protection against re-bleeding. In addition to the re-bleeding risk, there is a risk of overlooked DL in another part of the GI system. Risk of overlooked DL increases in the presence of ulcers. The demographic information, treatment methods, failed methods, re-bleeding rates, follow-up data obtained from the previous studies are given in Table 2.

There are several treatment algorithms for DLs that are presented in the literature. As a general rule, initial resuscitation is universal for all types of bleeding from DLs. Differences depend on the availability and experience at the centers. Although minimally invasive techniques are currently gaining more importance, the best way is to treat the patient with DL bleeding should depend on the availabilities of the health center. Early intervention and a cautious approach is the most important step in DL diagnosis and treatment.

Conclusion
DLs are rare, but it is important to be aware of this condition because DLs can cause massive hemorrhaging that can be life threatening. The location of the lesion and the hemodynamic status of the patient are the most important factors in the diagnosis and treatment modalities. It is preferable to use endoscopic methods for diagnosis in patients who are hemodynamically stable. In addition to diagnosis, multiple endoscopic treatments can be performed concurrently or in combination. To decrease the number of endoscopic sessions needed to achieve hemostasis, it is important to select the best treatment options and be aware of possible DLs when evaluating a patient who has GI bleeding. Surgical treatment is advantageous with low risk of re-bleeding, and surgery should be selected when patients are hemodynamically unstable and when other methods have failed. Additionally, the facilities and surgical experience of the centers to which patients are referred have a clear role in determining the selected treatments in their approach to treating DL patients. In conclusion, we recommend that clinicians develop a patient-specific strategy for DL patients, which includes consideration of the facilities at the centers in which they work.

Disclosure
The authors report no conflicts of interest in this work.

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