Preoperative Superselective Mesenteric Angiography and Methylene Blue Injection for Localization of Obscure Gastrointestinal Bleeding

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Despite a wide range of investigational techniques to aid localizing sites of gastrointestinal (GI) bleeding, approximately 5% of patients continue to bleed from an unidentified source.1,2 In these cases, the small bowel usually becomes the focus of inquiry when the site of bleeding is not apparent following standard investigations. Obscure GI bleeding (OGIB) has been reported to be of small-bowel origin in 45% to 75% of patients.2-5 We describe 4 patients with OGIB who had preoperative localization of the bleeding site with superselective mesenteric angiography, which was confirmed by the use of intraoperative methylene blue injection. This novel technique allowed us to identify the abnormal pathology, and, consequently, resection of the implicated segment of small bowel was performed without any postoperative complications. Final histology showed that 2 patients had arteriovenous malformations: one had a benign hemangioma of the small bowel, and the other had chronic ischemic ulceration in the ileum. Superselective mesenteric angiography combined with intraoperative localization with methylene blue is an important and innovative technique in the management of patients with unclear sources of gastrointestinal bleeding and allows for effective hemorrhage control with a focused and therefore limited bowel resection.

Methods

Of the emergency patients referred for OGIB, 4 were managed in our surgical department with this novel technique because the standard investigations failed to reveal the source of hemorrhage, and the transfusion requirements continued. All patients had an upper and lower GI endoscopy, followed by a capsule endoscopy. In addition, 2 patients had a push enteroscopy. Because the source of bleeding was not identified using these procedures, a multidisciplinary decision for SSMA followed by surgery was made after a discussion between the gastroenterologist, the interventional radiologist, and the surgeon. All patients were prepared to go straight to the operating room after undergoing visceral angiography. A superior mesenteric angiogram was performed in the standard fashion via the femoral artery route. Following localization of the abnormality, a guide wire was passed, and a 3F highly selective catheter was passed in the distal arterial branch over the guide wire. The catheter was secured and left indwelling, while the patient was transferred to the operating room. One senior surgeon (L.R.J.) performed all of the procedures. At exploratory laparotomy, 0.5 mL of methylene blue dye (10 mg/mL) was injected into the catheter, which resulted in immediate demarcation of the small intestine over a length of about 20 cm with brilliant blue color (Video). The delineated segment of bowel was resected in each patient with adequate margins, followed by primary anastomosis.

Results

Patient characteristics and outcomes can be seen in our Table. For patients with an active OGIB detected on a visceral angiographic scan, the SSMA technique had a 100% success rate. During this period, a total of 6 patients underwent visceral angiography, and of these 6 patients, 3 had active bleeding; therefore, SSMA was performed. The other 3 patients had no initial active extravasation and did not need subsequent intervention because their bleeding stopped spontaneously.

Localizing obscure gastrointestinal bleeding can be a clinical challenge, despite the availability of various endoscopic, imaging, and visceral angiographic techniques. We reviewed the management of patients presenting with obscure gastrointestinal bleeding during the period from 2005 to 2011. Four patients had preoperative localization of the bleeding site with superselective mesenteric angiography, which was confirmed by the use of intraoperative methylene blue injection. This novel technique allowed us to identify the abnormal pathology, and, consequently, resection of the implicated segment of small bowel was performed without any postoperative complications. Final histology showed that 2 patients had arteriovenous malformations: one had a benign hemangioma of the small bowel, and the other had chronic ischemic ulceration in the ileum. Superselective mesenteric angiography combined with intraoperative localization with methylene blue is an important and innovative technique in the management of patients with unclear sources of gastrointestinal bleeding and allows for effective hemorrhage control with a focused and therefore limited bowel resection.

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However, 1 patient rebled and had to undergo an SSMA following another visceral angiography that detected OGIB (Figure). Intraoperatively, there were no palpable lesions found in 3 patients, while 1 patient had a small nodule in the ileum that could have been easily overlooked. Three patients underwent a resection of the ileum, and 1 patient underwent a segmental resection of the jejunum. The median length of resection was 10 cm (range, 9-12 cm). Final histology showed arteriovenous malformations in 2 patients, a benign hemangioma in 1 patient, and a nonspecific chronic ischemic ulceration in 1 patient. All patients had an uneventful postoperative recovery. At median follow-up of 18 months (range, 10-24 months), no patients experienced rebleeding.

### Table. Clinical Characteristics and Outcomes of Patients Undergoing SSMA

<table>
<thead>
<tr>
<th>Patient No./Sex/Age, y</th>
<th>Presentation</th>
<th>Investigations Performed</th>
<th>Time Trends in Hb,(^a) g/dL</th>
<th>From Presentation to SSMA, d</th>
<th>In Angiography Suite, h</th>
<th>Blood Products,(^b) Units</th>
<th>Operative Finding</th>
<th>Surgical Resection</th>
<th>Histology</th>
<th>Time in OR, h</th>
<th>Complications</th>
<th>LOS,(^c) d</th>
<th>Outcome(^d)</th>
</tr>
</thead>
<tbody>
<tr>
<td>1/F/53 Melena, anemia</td>
<td>EGD, colonoscopy, CE, PE, angiography</td>
<td>6.7, 8.0, 10.4</td>
<td>10</td>
<td>2.5</td>
<td>12, 2</td>
<td>No palpable lesion</td>
<td>Ileum (10 cm)</td>
<td>AVM</td>
<td>3</td>
<td>None</td>
<td>7</td>
<td>No rebleeding at 24 mo</td>
<td></td>
</tr>
<tr>
<td>2/M/74 Melena, anemia</td>
<td>EGD, colonoscopy, CE, angiography</td>
<td>8.2, 7.6, 11.5</td>
<td>8</td>
<td>2</td>
<td>8, 0</td>
<td>No palpable lesion</td>
<td>Jejunum (10 cm)</td>
<td>AVM</td>
<td>2.5</td>
<td>Postoperative ileus</td>
<td>12</td>
<td>No rebleeding at 18 mo</td>
<td></td>
</tr>
<tr>
<td>3/F/55 Rectal bleeding</td>
<td>EGD, colonoscopy, CE, PE, angiography</td>
<td>5.8, 7.4, 10.9</td>
<td>12</td>
<td>3</td>
<td>18, 3</td>
<td>No palpable lesion</td>
<td>Ileum (12 cm)</td>
<td>Benign hemangioma</td>
<td>2.5</td>
<td>Thrombophlebitis</td>
<td>9</td>
<td>No rebleeding at 18 mo</td>
<td></td>
</tr>
<tr>
<td>4/M/53 Anemia</td>
<td>EGD, CE, angiography</td>
<td>7.2, 7.8, 11.6</td>
<td>6</td>
<td>2</td>
<td>9, 0</td>
<td>Small nodule</td>
<td>Ileum (9 cm)</td>
<td>Chronic ischemic ulceration</td>
<td>3.5</td>
<td>None</td>
<td>9</td>
<td>No rebleeding at 10 mo</td>
<td></td>
</tr>
</tbody>
</table>

Abbreviations: AVM, arteriovenous malformation; CE, capsule endoscopy; EGD, esophagogastroduodenoscopy; Hb, hemoglobin; LOS, length of hospital stay; OR, operating room; PE, push enteroscopy; SSMA, superselective mesenteric angiography.

\(\text{a}\) For each patient, the 3 values represent the Hb level at day of hospital admission, day of SSMA and surgery, and day of hospital discharge, respectively.

\(\text{b}\) For each patient, the 2 values represent red blood cells and fresh frozen plasma, respectively.

\(\text{c}\) After SSMA and surgery.

\(\text{d}\) Rebleeding was defined as further evidence of gastrointestinal hemorrhage in the form of melena, rectal bleeding, or unexplained anemia.
Localization of Obscure Gastrointestinal Bleeding

Discussion

Although uncommon, the management of OGIB can often be difficult and frustrating, especially if the bleeding originates from the small bowel. Angiodysplasia is the most common cause of acute or chronic hemorrhage from the small bowel, and these lesions are difficult to diagnose and locate during laparotomy.8 When patients present with exsanguinating GI bleeding, emergency surgery may be the only lifesaving possibility, after all nonoperative interventions have been tried (eTable in Supplement). Intraoperatively, lesions may be identified by visual inspection, illumination, or palpation. However, up to 50% of small-bowel lesions can be missed during laparotomy.9 An aggressive approach to preoperative localization of the bleeding source may prevent the high morbidity and mortality rates of blind surgical resection performed in desperation.10 Indeed, the lowest rebleeding rates after bowel resection for angiodysplasia have been reported in these situations.11-13 Fogler et al14 first supported the injection of methylene blue through a catheter placed in the superior mesenteric artery during laparotomy and localized a jejunal arteriovenous malformation on the basis of more rapid clearance of the dye from the lesion. This method had a distinct disadvantage in that an extensive dissection of the superior mesenteric artery was required. Improving on that, Athanasoulis and colleagues15 described an intraoperative localization procedure by injecting methylene blue through a superselective prepositioned angiographic catheter. This typically results in the staining of a segment of bowel 10 to 40 cm in length, which allowed for the operating surgeon to more accurately define the affected segment, resulting in more specific and successful surgery.16-18 A similar technique has also been used to identify the source of OGIB in infants.19

More recently, Stauffer et al,20 García-Barasa et al,21 and Stanojević et al22 have also successfully used this technique. Localizing OGIB intraoperatively by fluorescein injection through a superselectively placed catheter has also been reported.23 However, orange fluorescein dye is more tedious to use and requires the use of a UV Wood’s lamp intraoperatively. The successful use of provocative angiography (ie, the administration of 5000 U of heparin systemically and 10 mg of tissue plasminogen activator injected during angiography directly into the superior mesenteric artery) followed by superselective injection of methylene blue has also been reported.24 Of note, one of the limitations of our technique is that the hemorrhage has to be substantial and active at the time of the study in order to localize the source. Furthermore, when angiography can accurately locate an abnormality, the subsequent intraoperative localization of the bleeding point can still be a challenge. Superselective angiographic catheters have enabled interventional radiologists to cannulate distal visceral vessels and more accurately pinpoint the site of a lesion or bleeding. Intraoperative methylene blue injection causes a local, immediate, and dramatic demarcation of the small bowel that allows the suspected area to be well defined.25 The recommended maximum intravascular dose is 1 to 2 mg/kg,17 and each injection uses 0.5 mL of methylene blue (5 mg); thus, multiple injections can be made without reaching the maximum dose. Our small series of 4 cases exemplifies the problems that are commonly encountered in identifying the source of OGIB. Even though our cases all had pathology in the small bowel, we believe that this method could also be applied to cases with obscure colonic bleeding. Because surgery still offers the best chance for cure, we believe that SSMA combined with intraoperative localization with methylene blue dye plays an important role in the diagnostic and therapeutic workup of these challenging cases as an adjunct to standard endoscopic and radiological investigations. Larger prospective studies assessing the use of SSMA are warranted.