Introduction

Strictures can occur at any level of the gastrointestinal tract (GIT) from the esophagus to the colon. The causes of GIT strictures are numerous, such as reflux disease, congenital webs, caustic ingestion, malignancy, anastomotic, radiation, Schatzki’s rings, pyloric stenosis, nonsteroidal anti-inflammatory drug (NSAID) use, diverticulitis, or inflammatory bowel disease. In addition, these strictures are difficult to treat at times as each intervention may lead to further scarring and worsening stricture formation. As one may expect with such various etiologies and locations, there are many modalities of treatment. Historically, the first treatments involved dilation with fixed diameter bougies. As anesthesia and the field of surgery progressed, more invasive surgeries were developed that would allow for the resection of the diseased area. More recently, adjunctive endoscopic therapies such as stent placement, intralesional injections and electrosurgery have evolved to help solve this complex disease process.

Dilation

The earliest treatments were focused on the esophagus as it was the most accessible of areas. In the mid-1600s, Dr. Thomas Willis described using a thick sponge on the end of a whalebone bougie and in the 1700s, some doctors in Europe were recording the use of ivory bougies to dilate esophageal strictures (1). Over the past 150 years, various types of dilators have been used. Initially these were placed blindly, but fluoroscopic guidance has become a useful adjunct to aid with visualization during dilation. As the field of surgery progressed, retrograde dilators were able to be used by having the patient swallow a string which was then attached to a dilator through a gastrostomy site and pulled in a retrograde fashion through the esophagus. Another technique that has been used was using string guidance, whereby a patient swallowed a weighted string and after this passed into the distal esophagus past the stricture, GIT dilators were then passed over the taut string.

The most common blind bougie dilators in use today are
the Hurst and Maloney dilators (Figure 1). Both are filled with tungsten and push type dilators that exert a shearing and radial force as they are pushed through a stricture. One point of differentiation with these two dilator types is that the Maloney dilators have a tapered type tip while the Hurst dilators have a blunt style tip. Another option that allows for slightly more control are Savary-Gilliard, American Dilation System and Tucker dilators that have an internal channel that allows for wire guidance. The Savary-Gilliard and American Dilation System dilators are push type dilators while the Tucker dilators are of the pull type. Tucker dilators have a loop on both tapered ends. They are pulled both antegrade and retrograde across a stricture. The Savary-Gilliard dilators have a radiopaque marking at the end of the taper point that is the maximum diameter whereas the American Dilation System dilators are totally radiopaque. Finally, anorectal Hegar dilators are stainless steel dilators with rounded ends. They are short and tend to be used for anorectal strictures. These dilators all have fixed maximum diameter and exert both radial and longitudinal shearing forces on the stricture during dilation.

In 1971, Lilly et al. reported the first endoscopic guided dilation which allowed a whole new avenue of therapies and began the use of balloon expandable dilators (2). These dilators come in various sizes ranging from 5 to 20 mm in diameter and exert only a radial force. By using the endoscope for guidance, it is possible to access strictures throughout the GIT as compared to the prior available dilators. They are designed to pass through the therapeutic channel of an endoscope with or without wire assistance. If inflated with radiopaque contrast, one is able to observe the dilation throughout the length of the stricture (Figure 2). The balloons are single use devices which can add to the cost of the procedure compared to bougie dilators. A handheld device to inject liquid is used to inflate the balloon, which then allows the proceduralist to monitor the pressure used during the dilation in addition to observing it through the endoscope. Over the years, a few trials have investigated comparing fixed diameter bougie dilators with endoscopic guided balloon dilators, mostly in the treatment of esophageal strictures as these are the most accessible to both treatment modalities. On a recent meta-analysis, no difference was noted in symptomatic relief, recurrence at one year, bleeding or perforation. However, patients who had balloon dilation did experience less severe post procedure pain (4).

**Adjunct therapy**

Although the majority of strictures can be effectively managed with endoscopic dilatation, additional measures can also be considered in the management of strictures depending on the etiology. Stent placement is one of the most common adjunctive modalities to maintain lumen...
patency, particularly with malignant strictures. Stents act by exerting radial force on the stricture; there are a variety of stents available depending on the location, length, material and diameter (Figure 3) (5).

Uncovered stents are generally used in patients with poor survival and in areas with expected high likelihood of stent migration such as the duodenum and colon because of its benefit of flexibility to exert high pressure on angulated strictures in these regions (6). However, due to the risk of tumor and tissue in-growth with uncovered metal stents that can lead to re-obstruction and symptom recurrence, the majority of stents used for malignant strictures are either partially-covered or fully-covered metal designs, which have both been found to be equally effective and safe (7). Although fully-covered stents are more resistant to tumor and tissue in-growth, one of their disadvantages is that they are more prone to stent migration which can require re-intervention. Strategies to reduce the risk of stent migration include the use of large-sized stents and anchoring of the stents using clips at the stent margin or endoscopic sutures (8). Partially-covered stents do offer some of the advantages of both uncovered and fully-covered stents; however, in benign pathologies, partially covered stents are not recommended due to the proliferation of granulation tissue through the proximal and distal uncovered stent edges which can make removal difficult. As such, fully-covered metal or self-expanding plastic stents are commonly used for benign strictures as a temporary treatment strategy which has the benefit that they are more easily removed compared to uncovered stents that can be more challenging to remove due to tissue ingrowth. To prevent excessive tissue in- or overgrowth, it is generally recommended that stents be removed by 6–10 weeks after treatment (9). Techniques for stent removal include using endoscopic rat tooth forceps to pull the purse-string suture at the proximal stent edge, double forcep graspers through a double-channel endoscope, stent-in-stent technique of temporary placement of a new stent within the first stent to facilitate removal and stent inversion of grasping the distal end of the stent and inverting it through itself. There is new evidence supporting the use of biodegradable stents which are typically made of polydioxanone that would not require removal; a recent multi-center randomized control trial found that biodegradable stent placement was associated with a significantly longer median time to first dilation with improved symptom control and higher level of activity compared to dilation alone with no difference in adverse events (10). However, further studies are still required to define the role of biodegradable stents in the treatment of benign strictures. Regardless of stent type, there was no significant difference found between stent types in the treatment of benign strictures, with an approximate success rate of 40% in benign esophageal strictures (11). The role of endoscopic stent placement will also be further discussed in another article in this series.

Endoscopic intralesional steroid injections have also been considered as an adjunctive therapy either before or after dilation and are typically used for refractory benign or anastomotic strictures. It is believed that intralesional steroids inhibit stricture formation by interfering with collagen synthesis, fibrosis and chronic scarring processes to help augment the effects of dilation but the mechanism of action is not entirely clear (12). The steroid utilized is typically triamcinolone acetate or acetonide in a concentration of 10–40 mg/mL injected using a 21 to 23 gauge 5 mm long sclerotherapy needle in 0.5 mL aliquots circumferentially in four quadrants at the proximal margin of the stricture as well as in the strictured segment when possible which should be difficult to inject if in the correct position (13). Although a few small studies demonstrated symptom improvement and an overall reduction in the need for repeat dilatations for benign esophageal strictures, these were not found to be statistically significant in a
more recent meta-analysis. However, it was found to be consistent across all studies that there was a significant increase in the interval between dilations indicating some efficacy of intraluminal steroids when used in combination with endoscopic dilation (14). The efficacy of intraluminal steroids in addition to endoscopic balloon dilation was also found to be replicated with anastomotic strictures following esophagectomy in which a small randomized control trial demonstrated a significant reduction in the total number of dilations required as well as a significant improvement in re-stricture free survival (15).

With a similar logic as intraluminal steroid injections that help counter collagen synthesis, fibrosis and scarring, people have investigated topical application of mitomycin C to strictures. In an animal study, mitomycin C in a dose dependent manner showed significant ability to prevent stricture formation after caustic esophageal injury with fairly good results which was then replicated in a case series. A double-blind, randomized, placebo controlled trial showed 80% resolution of esophageal strictures with topical mitomycin C treatment compared to only 35% in the placebo group as well as less total number of dilation sessions in the treatment group (16). Mitomycin C has also been injected intra-lesionally into the submucosal space in complex recurrent esophageal strictures, in which a study found that there was a significant improvement and resolution of dysphagia symptoms in the group treated with mitomycin C compared to the placebo group with no adverse effects (17). In addition, mitomycin C was found to be effective in stricture prevention in a porcine model (53.6% low dose, 35% high dose); however, in this trial, the esophageal wall treated with high-dose mitomycin C appeared to be necrotic and eventually led to perforation. In models treated with a low dose of mitomycin C, the esophageal wall appeared re-epithelialized and healthy (18).

Another adjunctive measure to endoscopic dilations used for benign refractory and anastomotic strictures is electroincisional therapy in which electrocautery can be combined with argon plasma beam coagulation, needle-knife or endoscopic scissor techniques to incise the tissue of the stricture. The most common technique is the radial incision and cutting technique, in which the stricture is incised under direct endoscopic vison in a radial fashion parallel to the longitudinal axis to remove the rim of the stenosis (Figure 4) (19). The procedure is typically terminated once the scope can easily traverse the previously strictured segment (20). Incisional therapy can also be used in conjunction with dilation or steroid injections to reduce the need for repeated intervention. One randomized controlled trial found that electroincisional therapy was comparable in efficacy in terms of the total number of dilations and success rate when compared with Savary dilations with no increase in adverse events (19).

Current evidence also suggests that incisional therapy is also most effective in the treatment of refractory benign or anastomotic strictures that have a relatively short stenosis less than 1cm in length with a success rate of approximately 80.6% and rate of recurrence of approximately 4.8% (21). As such, electroincisional therapy can be considered a safe and effective alternative method for the treatment of refractory strictures for whom multiple re-interventions have failed.
Complications

Given that the goal of dilation is to disrupt the stricture to enlarge the lumen, mucosal tears are expected. However, significant complications can occur which can include bleeding, perforation, bacteremia and aspiration which is mostly with upper GIT strictures and also related to sedation given for the procedure. Minor bleeding following dilation, stent insertion or electroincisional therapy is relatively common and typically self-limiting not requiring any intervention as fibrotic strictures are relatively avascular; however, significant bleeding from these interventions occurs at a rate between 0.1–0.4% (19,22). With this risk in mind, the American Society of Gastrointestinal Endoscopy (ASGE) came out with guidelines to help address the added bleeding risk of therapeutic endoscopy on patients who are on anti-coagulation or antithrombotics. For high-risk procedures such as dilation or incisional therapies, anti-coagulation and thienopyridines should be held for the appropriate time frame while non-steroidal anti-inflammatory drugs, aspirin and DVT prophylaxis should be continued even in high risk procedures. Controversially, endoluminal stent placement is categorized as low risk and as such, it is recommended to continue anti-coagulation in this setting (23). Perforation is the most clinically significant complication and can occur either from complete transmural disruption or creation of a false tract (24). It is estimated that the overall risk of perforation ranges from 0.1% to 0.6% (25). Factors that are associated with an increased risk of perforation include malignancy, severe inflammation, radiation or caustic-induced strictures as well as operator inexperience. Features of strictures that also increase the risk of perforation include long-segment, significant luminal narrowing or angulated strictures (26). Bacteremia is also a known complication from endoscopic dilation; however, it is rarely clinically significant as it is equivalent with brushing and flossing and the current ASGE guidelines do not recommend routine antibiotic prophylaxis at the time of endoscopic intervention for the management of strictures (27). Ultimately, endoscopic management of strictures is generally safe, effective and has a favorable risk-benefit profile. Additional studies are required to better define the long-term efficacy of endoscopic options such as intralesional steroid or mitomycin C injections, biodegradable stents and electroincisional therapy.

Acknowledgments

The authors would like to thank Ryan Juza and Jeffery Marks for allowing us to participate in this special issue on surgical endoscopy.

Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

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doi: 10.21037/ales.2019.06.06

Cite this article as: Johnson R, Fung EC. Dilation and adjunct therapy for strictures. Ann Laparosc Endosc Surg 2019;4:63.