Percutaneous Management of Benign and Postoperative Biliary Strictures

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Most benign biliary strictures (BBSs) are most commonly iatrogenic in nature and are typically the result of surgical injury. Postoperative strictures can be broadly divided into anastomotic and nonanastomotic strictures according to their anatomic locations. Anastomotic strictures typically develop in patients who have undergone previous hepatobiliary surgery or liver transplantation. Liver transplant patients have a relatively higher incidence of biliary anastomotic stricture than those with other types of abdominal surgeries.1

Common etiologies for nonanastomotic BBS include cholangitis, ischemic injury during liver transplantation, and cholelithiasis. Clinical manifestations of BBS include jaundice, fever, elevated levels of serum alkaline phosphatase, and bilirubin. Endoscopic retrograde cholangiopancreatography (ERCP) is recognized as the first-line treatment for BBS2; however, ERCP is not a viable option for patients with Roux-en-Y reconstruction or those with esophageal or upper gastrointestinal obstruction. In these situations, alternative approaches to access the bile ducts are necessary for diagnosis and treatment.3

Interventional radiologists play a vital role in the management of BBS by providing image-guided percutaneous transhepatic cholangiography (PTC), catheter insertion, balloon dilation, and stent placement. The purpose of this review is to summarize percutaneous techniques, short-term and long-term therapeutic effects, complications, and novel techniques for management of BBS.

Abstract

Biliary strictures are an uncommon yet challenging clinical problem. They are often iatrogenic in nature, usually the consequence of hepatobiliary surgery. Although the etiology may be benign, the clinical consequences, if unrecognized, can progress to ascending cholangitis, sepsis, and hepatic abscesses. Optimal treatment requires expertise and input from multiple specialties, including gastroenterology, surgery, infectious disease, and interventional radiology. Currently, available interventional techniques play a critical role in the management of patients with benign biliary strictures.

Keywords

► benign biliary stricture
► percutaneous intervention
► biliary obstruction
► biliary stents

Percutaneous Transhepatic Cholangiography

The purpose of PTC is to gain access and evaluate the intrahepatic bile ducts by means of contrast injection. Successful percutaneous puncture into a dilated intrahepatic biliary tree with a fine needle is essential for access to biliary system and subsequent diagnosis and treatment. Needle puncture of dilated bile ducts can be performed using under fluoroscopy, ultrasound, or computed tomography (CT) guidance. Accessing the biliary tree can be challenging in postoperative patients with altered ductal anatomy or reduced liver volume.4 Auxiliary techniques to improve access to the biliary tree in cases of altered anatomy have been described. In 1967, opacification of the biliary tree via puncture of the gallbladder was shown to facilitate secondary biliary
puncture. A randomized study comparing CT fluoroscopy with fluoroscopy alone found that CT fluoroscopy combined with fluoroscopy led to higher successful rates of first puncture, fewer punctures, and shorter procedure times. At times, it may be necessary to access a nondilated biliary system. In these situations, access can often be achieved by directing needle punctures in the vicinity of peripheral branches of the portal vein. In a study evaluating this technique, 35 patients with biliary strictures with nondilated bile ducts, including 24 cases of benign biliary anastomotic strictures, successful access to the biliary tree was achieved in 94.3% of cases. This technique is very helpful for the postliver transplantation patients when early biliary strictures are suspected.

PTC can accurately depict the location and characteristics of biliary strictures in most patients. The appearances and features of BBS in cholangiography can vary depending on the etiology. Most biliary strictures are unifocal and are often iatrogenic (liver transplantation, partial hepatectomy, and bile duct injury). Noniatrogenic causes such as stone disease, pancreatitis, trauma, radiation therapy, portal biliopathy, and idiopathic papillary stenosis can also lead to unifocal strictures. The reasons for multifocal or diffuse bile duct injury). Noniatrogenic causes such as stone disease, pancreatitis, trauma, radiation therapy, portal biliopathy, and idiopathic papillary stenosis can also lead to unifocal strictures. The reasons for multifocal or diffuse bile duct strictures include ischemia (hepatic artery occlusion after liver transplantation) and cholangitis (sclerosing cholangitis, pyogenic cholangitis, tuberculosis, human immunodeficiency virus, parasitosis, autoimmune disease, and immunoglobulin G4-related cholangiopathy). The goals of PTC are to depict the levels of obstruction, evaluate for bile duct stones, define etiologies of cholangitis, and demonstrate bile duct leak. Additional maneuvers such as catheter drainage, balloon dilation, and stent placement can be performed through PTC access.

**Percutaneous Transhepatic Catheter Drainage**

The goal of catheter drainage of the biliary is to relieve jaundice, control infection, and improve nutritional status, which are all essential for patients in the perioperative period. The decision regarding whether to do more invasive manipulations at the time of initial drainage depends on technical challenges or complications during bile duct catheterization. When cholangitis, bleeding, or infection complicate biliary catheterization, or if the puncture has been technically challenging and time consuming, an external biliary drainage catheter should be placed without further attempts to cross the stricture. Alternatively, if PTC goes smoothly, attempts to cross biliary strictures with guide wires are reasonable. It may not always be easy to cross the strictures by using a conventional 0.035” guidewire. Fidelman introduced a Lunderquist-Ring Torque guidewire with a “stick” formation at the distal end and a “handle” formed on the proximal end to allow torqueing, which is useful for crossing strictures. In circumstances of complete biliary occlusion, needle–knife fistulotomy may be considered to re-establish communication between the biliary tree and bowel. Usually, an external biliary drainage protocol with or without balloon dilation will be undertaken to achieve restoration of biliary patency.

**Balloon Dilation**

Selection of balloon size should be estimated based on the cholangiogram. Initially, a balloon size slightly less than or equal to bile duct diameter adjacent to the stricture is recommended for initial dilatation to minimize the risk of anastomotic disruption and bile leak. If necessary, larger diameter balloons can be used for subsequent dilations with balloon sizes up to 25 to 30% larger than the estimated diameter of the duct being dilated. Common bile duct
strictures can be safely dilated to 10 to 12 mm in an adult. Smaller balloons (4–8 mm) may be necessary for lobar ducts and for the biliary strictures in children.\textsuperscript{9,23} Time intervals between balloon dilations range from 1 to 2 weeks to every 3 months to establish and maintain duct patency.\textsuperscript{17,18} Angioplasty balloon catheters should always be inflated gradually until the site of narrowing, commonly referred as the “waist” is eliminated (Fig. 1). Satisfactory dilation is marked by disappearance of the “waist” and restoration of normal bile flow.\textsuperscript{24}

**Fig. 1** (A) Color Doppler ultrasound image of a 63-year-old patient s/p complicated cholecystectomy that demonstrates dilated intrahepatic bile duct (white arrow). (B) Axial contrast material-enhanced CT image that demonstrates dilated biliary branches in left and right hepatic lobes (white arrows). (C) MRCP that demonstrates bilioenteric anastomosis stenosis (white arrow). (D) Percutaneous transhepatic cholangiogram that demonstrates dilated bile ducts and anastomotic stricture. (E) Balloon dilation of the anastomotic stricture from a percutaneous right duct approach. “Waist” (white arrow) indicates site of stricture. (F) Balloon dilation of left biliary stricture that demonstrates “waist” (white arrow). (G) Postprocedure cholangiogram that demonstrates 12F catheters placed across the bilioenteric anastomosis via left and right ducts. (H) Transhepatic cholangiogram 3 months after balloon dilation and continuous drainage shows resolution of anastomotic stricture and widely patent anastomosis (white arrow). The catheters were removed after cholangiography. (I) Axial contrast material-enhanced CT 6 months after catheter removal shows decompressed bile ducts. CT, computed tomography; MRCP, magnetic resonance cholangiopancreatography.
The use of cutting balloons has been described for refractory biliary strictures, especially those unresponsive to conventional balloon dilatation. Cutting balloons are designed with four longitudinally mounted, 1-cm long microsurgical blades that are exposed during inflation. Several reports suggest that cutting balloon is associated with high success rates and few complications.25–28 Mukund et al reported a group of eight cases of biliary strictures treated with cutting balloon with up to 14 months of follow-up free of recurrent biliary obstruction and cholangitis.25 Further randomized contrast clinical trials are necessary to verify its safety and efficacy.

**Long-Term Results of Catheterization and Balloon Dilation**

From 2007 to present, eight studies with the mean follow-up time of 34 months to 5.2 years have evaluated the long-term results of balloon dilatation.10–12,14–18 Most have showed 1-year patency rates of 94 to 100%, while 3-year patency rates range from 71 to 96%.10–12,15,16,18 The 5- and 10-year patency rates were 74 to 88 and 67 to 72%, respectively, in two recent studies.12,14 However, a retrospective study with 25-year results showed a lower long-term patency of 52, 49, and 41% in 5, 10, and 25 years, respectively.17 The long-term results of catheterization and balloon dilation are listed in Table 1.

Kucukay et al found that there was no difference in long-term patency between patients who underwent successful one-time balloon dilation versus those with successful initial dilatation and two or more subsequent dilations (85.4 vs. 87.9%). As a result, repeated balloon dilation is not recommended when the first procedure is successful.24 Furthermore, Janssen et al found that restenosis and treatment failure occurred more often in patients who underwent multiple dilatations. They found that factors that led clinically relevant restenosis were the number of strictures and the number of treatments.12 Between strictures at anastomotic and non-anastomotic sites, Cantwell et al revealed that no significant difference was found in the rate of clinically significant restenosis after the first treatment.17 Regarding the patients with orthotopic liver transplantation (OLT), the risk of stricture recurrence for those with no history of OLT was found to be lower than that for patients with a history of OLT.14

**Biliary Stents**

Self-expandable metallic stent (SEMS) provides an optional solution for refractory or recurrent BBS, especially when patients are unsuitable or unwilling to undergo surgery.29,30 However, the use of SEMS for BBS remains controversial at present because of the high rate of stent obstruction secondary to epithelial hyperplasia.31 Furthermore, stent removal
<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>Cases</th>
<th>Catheter size (Fr)</th>
<th>Catheter Size (Fr)</th>
<th>Catheter change interval</th>
<th>Drainage duration</th>
<th>Balloon diameter (mm)</th>
<th>Follow-up time</th>
<th>Long-term patency rate (y) (%)</th>
</tr>
</thead>
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<tr>
<td>DePietro et al</td>
<td>2015</td>
<td>71</td>
<td>12–14</td>
<td>16–18</td>
<td>3 mo</td>
<td>74 (4–367) d</td>
<td>8</td>
<td>4.7 (0–12) y</td>
<td>1 2 5 10</td>
</tr>
<tr>
<td>Janssen et al</td>
<td>2014</td>
<td>98</td>
<td>8.5</td>
<td>12</td>
<td>3 wk</td>
<td>14 (2–69) wk</td>
<td>4–10</td>
<td>35 (0–146) mo</td>
<td>1 2 5 10</td>
</tr>
<tr>
<td>Bonnel and Fingerhut</td>
<td>2012</td>
<td>110</td>
<td>10</td>
<td>NA</td>
<td>NA</td>
<td>8.5 (4.5–45) mo</td>
<td>10</td>
<td>59 (0.5–278) mo</td>
<td>1 2 3 7.4</td>
</tr>
<tr>
<td>Gwon et al</td>
<td>2011</td>
<td>79</td>
<td>8.5–10</td>
<td>14 + 8.5 (dual catheter)</td>
<td>2 mo</td>
<td>6.5 (5.5–14.2) mo</td>
<td>6–8</td>
<td>34.5 (6.1–26.2) mo</td>
<td>1 2 3</td>
</tr>
<tr>
<td>Weber et al</td>
<td>2009</td>
<td>27</td>
<td>8.5</td>
<td>16</td>
<td>NA</td>
<td>19.9 mo</td>
<td>NA</td>
<td>53.7 mo</td>
<td>100% during follow-up</td>
</tr>
<tr>
<td>Glas et al</td>
<td>2008</td>
<td>38</td>
<td>15</td>
<td>NA</td>
<td>3 m</td>
<td>346 (126–488) d</td>
<td>7–14</td>
<td>34 (11.3–65.2) mo</td>
<td>1 2 3</td>
</tr>
<tr>
<td>Cantwell et al</td>
<td>2008</td>
<td>85</td>
<td>7–10</td>
<td>10–12</td>
<td>NA</td>
<td>28–42 d</td>
<td>10–12</td>
<td>Unknown</td>
<td>5 10 15 20 25</td>
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<td>Köcher et al</td>
<td>2007</td>
<td>21</td>
<td>12</td>
<td>NA</td>
<td>NA</td>
<td>10.3 (3–11) m</td>
<td>10–12</td>
<td>5.2 (16 mo–11 y)</td>
<td>1 2 3</td>
</tr>
</tbody>
</table>

Abbreviation: NA, nonapplicable.
may be difficult or impossible once they become embedded in the wall due to epithelial hyperplasia.\textsuperscript{32}

The introduction of fully covered SEMS (FCSEMS) offers solutions to overcome these limitations. Covered metal stents have a metallic skeleton covered by a biocompatible, layer of synthetic material which is resistant to the effects of bile, gastric, and pancreatic secretions. The covered layer prevents the stent from becoming embedded and makes stent very smooth and easy to remove.\textsuperscript{33,34} In addition, FCSEMS can be implanted and removed by either endoscopic or percutaneous methods. Gwon et al reported 68 cases of BBS treated with percutaneously placed FCSEMS and showed technical success of 98.5\% (67/68) for stent placement and 98.5\% (66/67) for stent removal.\textsuperscript{35} After a mean stent indwelling period of 5.8 months and mean follow-up period of 36 months, the primary patency rates at 1, 2, 3, 4, and 5 years reached 91, 89, 76, 68, and 68\%, respectively.\textsuperscript{35} Another function of FCSEMS is to block the bile leaks, especially for the bilioenteric anastomotic strictures combined with leakage. Gwon et al reported 100\% clinical success of 11 cases of postoperative bile leaks treated with retrieval FCSEMS.\textsuperscript{36} Recently, multicenter clinical trials showed that endoscopic placed FCSEMS was safe and effective for BBS.\textsuperscript{37,38} Although FCSEMSs are effective in minimizing endothelial overgrowth, their smooth surfaces may contribute to stent migration which is reportedly as high rate as 16.2\%.\textsuperscript{35}

The application of biodegradable stents in BBS may help overcome the migration problems associated with FCSEMS. The currently available biodegradable biliary stent is made of polydioxanone, which is proved by the Food and Drug Administration as a biodegradable material for clinical use. Delivery systems range in diameter from 11 to 15 Fr and the degradation time, which occurs by hydrolytic processes, is approximately 3 to 6 months.\textsuperscript{39,40} Recently, Mauri et al reported a multicenter clinical study of percutaneously implanted of biodegradable stents for 107 patients with refractory BBS. The technical success was 98\%. Stent migration was observed only in 2/107 cases (2\%). The stent recurrence rates at 1, 2, and 3 years were 7.2, 26.4, and 29.4\%, respectively. Although they reported no major complications, restenosis was found to be associated with subsequent cholangitis and biliary stones.\textsuperscript{41}

### Other Percutaneous Techniques for Management of Benign Biliary Strictures

#### Magnetic Compression Anastomosis

Magnetic compression anastomosis (MCA) is a technique whereby a pair of cylindrical samarium–cobalt magnets that measure 4 × 10 mm are used to treat high grade or complete biliary strictures. Two magnets are introduced by percutaneous and endoscopic access separately and approximate each other on different sides of the stricture. Compression by the approximated magnets caused ischemic necrosis of the tissue between the magnets, forming a fistula. Jang et al reported their clinical effectiveness treating both biliobiliary and bilioenteric anastomoses.\textsuperscript{42,43}

#### Percutaneous Transhepatic Cholangioscopy

Percutaneous transhepatic cholangioscopy (PTCS) offers the option of direct endoluminal visualization of the biliary tree for diagnostic and therapeutic interventions.\textsuperscript{44} Biliary stones are the most commonly encountered disorder with biliary strictures. Treatment of BBS can be difficult in the setting of choledocholithiasis. Percutaneous transhepatic endoscopic biliary holmium laser lithotripsy (PTBL) can be helpful in this setting. Rimon et al reported a group of patients with biliary stones who failed conventional treatment that were treated with PTBL. In 6/22 patients with stone disease in the setting of biliary strictures, PTBLs were able to achieve complete fragmentation of the stones with no major complication occurred.\textsuperscript{45}

#### Percutaneous Transjejunal Biliary Intervention

Percutaneous transjejunal biliary intervention (PTJBI) is an alternative technique for access to the biliary system to treat benign biliary diseases in patients with a Roux-en-Y hepaticojejunostomy. Fontein et al reported the largest series that evaluated efficacy of PTJBI for BBS and stone retrieval. In this procedure, the Roux loop is punctured under fluoroscopic guidance, using surgical clips as a landmark, and a guidewire advanced into the afferent limb. Then, an 8- to 10-Fr sheath is advanced over the guidewire into the afferent limb, thus providing access to biliary strictures and stones if a guidewire crosses the hepaticojejunal anastomosis.\textsuperscript{46} Mansueto et al treated a group of patients with hepaticojejunostomy dehiscence in which initial access to the biliary tree was achieved by PTJBI, followed by a rendezvous technique to establish percutaneous biliary drainage to the jejunum.\textsuperscript{47}

#### Complications

Hemorrhage and infection represent the most commonly encountered complications of percutaneous biliary interventions. An analysis of 419 cases of percutaneous transhepatic biliary drainages indicated an overall complication rate of 9.31\%, including bleeding (2.86\%), cholangitis (1.67\%), and sepsis (1.43\%).\textsuperscript{48} Transarterial embolization is an effective management for bleeding associated with active extravasation or pseudoaneurysm of the hepatic artery.\textsuperscript{13,17,49} Biliary leak and bilomas are less frequently encountered and can usually be managed by drainage.\textsuperscript{50}

Drainage-related complications are reported as high as 21.4\%, including occlusion, dislocation, and cholangitis.\textsuperscript{51} Stent-related complications include stent migration and perforation.\textsuperscript{52} Procedure-related mortality was reported as 2\%. Multivariate analysis revealed this was related to the preprocedure presence of ascites, high C-reactive protein, and a high white cell count.\textsuperscript{53}

#### Conclusion

Percutaneous transhepatic interventions play an important role in the treatment of benign or postoperative biliary strictures, especially for those that are inaccessible by endoscopic techniques. Treatment can be accomplished by one of

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several interventional techniques. Catheterization and balloon dilation are the primary recommended methods for treating BBS, with the satisfactory long-term results. Newer techniques, such as SEMSs, fully covered, and biodegradable stents are options when strictures are refractory treatment or fail to resolve by conventional methods. Innovative and novel techniques such as MCA, PTCS, and PTJBI may be helpful in the management of BBS. However large, multicenter, randomized contrast clinical trials are necessary to further evaluate their long-term efficacy and safety.

Conflict of Interest
None.

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