INTRODUCTION

Despite the steady improvements in surgical techniques and postoperative patient care over the past decade, postoperative bile leakage remains a complication of various hepatobiliary surgeries. The incidence of bile leakage after laparoscopic cholecystectomy was reported to be 1.1% to 4.0% [1-5]. In addition, bile leakage occurs following ~15% of liver resections [6-12]. The frequency of hepatobiliary surgery for neoplastic diseases has increased due to early screening; this has resulted in an increase in the number of cases of bile leakage after surgery. Therapeutic options for bile leakage include percutaneous drainage (PCD) of fluid collection (such
as bilomas), therapeutic biliary endoscopy, and surgery. Recently, biliary endoscopic procedures including biliary stenting and/or endoscopic sphincterotomy (EST) have become the first choice for diagnosis and treatment of postoperative bile leakage. The primary aim of this study was to evaluate the efficacy of endoscopic treatment of bile leakage after hepatobiliary surgery.

**METHODS**

A retrospective review of 20 patients referred for post-hepatobiliary surgery bile leakage from August 2009 to September 2014 was performed. The biliary tract and leakage site in each patient were visualized during endoscopic retrograde cholangiopancreatography (ERCP). The medical records, endoscopic, and radiologic findings were reviewed retrospectively. Three experienced pancreatobiliary endoscopists performed all ERCP procedures. ERCP was performed using a side-viewing duodenoscope (TJF-240, Olympus Optical Co., Tokyo, Japan). Wire-guided cannulation of the biliary tract was performed, and cholangiograms were obtained. Endoscopic therapy in all patients consisted of EST to reduce intraductal pressure and facilitate transpapillary bile flow. A plastic stent with a diameter of 7 or 10 Fr was inserted, as determined by the endoscopist. Patients were discharged after resolution of leakage-related clinical symptoms, and follow-up ERCP was performed at 4 weeks for cases of postcholecystectomy bile leakage and at 4 to 8 weeks for cases of major hepatectomy to confirm complete healing of the leak and to remove the biliary stent. Success of ERCP was determined primarily as a decrease in percutaneous bile drainage volume through a Jackson-Pratt bag. Success was confirmed when extravasation of contrast medium was not seen on follow-up ERCP. If bile leakage was detected at follow-up cholangiography, stent patency was maintained for up to 8 weeks.

The study protocol was approved by the Institutional Review Board of Konkuk University Medical Center, which confirmed that the study was in accordance with the ethical guidelines of the Helsinki Declaration.

**RESULTS**

A total of 20 patients (18 males, two females; mean age, 51.4 ± 14.1 years) with postoperative bile leakage were referred for endoscopic management (Table 1). The surgical department of our hospital referred 16 patients; the remaining four patients were transferred to other institutions for treatment.

Thirteen patients had undergone laparoscopic cholecystectomy (65%) because of cholecystitis, two patients were living-donor recipients and had undergone lobectomy (10%) because of liver transplantation, three had undergone segmentectomy (15%) because of hepatocellular carcinoma, one had undergone segmentectomy (5%) because of rectosigmoid carcinoma with liver metastasis,
and one had undergone partial resection (5%) because of rectosigmoid carcinoma with liver metastasis (Fig. 1).

The mean time from surgery to diagnosis of bile leakage was 9.0 ± 6.4 days (range, 1 to 22). Most bile leakages presented as percutaneous bile drainage through a Jackson-Pratt bag (75%), followed by abdominal pain (20%) and fever (5%). The mean interval from hepatobiliary surgery to referral for ERCP was 12.4 ± 8.5 days (range, 2 to 31). The mean interval was longer than that reported in a previous study [13-15]. The four patients referred for treatment from other institutions had longer intervals than those of the other patients.

The mean interval from endoscopic retrograde biliary drainage (ERBD) to stent removal was 77.6 ± 21.7 days. The stent was not removed in three patients. Two of these three patients were lost to follow-up and the other patient died because of cancer aggravation. Bile drainage through a Jackson-Pratt bag was stopped at a mean interval of 5 days (range, 3 to 12), and percutaneous catheters were able to be removed at a mean interval of 6 days (range, 5 to 16) after endoscopic procedures.

The sites of bile leaks were the cystic duct stump in 10 patients (50%), right/left intrahepatic ducts in five (25%; right, three; left, two), liver beds in three (15%; duct of Luschka, two; cystic duct orifice, one), common hepatic duct in one (5%), and common bile duct (CBD) in one (5%). Of the 20 patients with postoperative bile leakage, three cases were combined with bile duct stricture (15%); one patient had severe CBD obstruction, and the others had mild stricture. Bile leakage occurred together with CBD stones in five cases. After EST and controlled radial expansion wire-guided papillary balloon dilatation, stone removal was performed using an extraction basket. A plastic stent was then placed at the site of bile leakage.

Concerning endoscopic modalities, endoscopic therapy for bile leak was successful in 19 patients (95%). One patient developed microperforation during cannulaion, which was mild, and recovered without clinical sequelae. In one case, the amount of percutaneous bile drainage increased 3 days after the endoscopic procedure. Additional biliary stenting with sphincterotomy was performed, which stopped the bile leakage.

One patient was diagnosed with hepatocellular carcinoma and underwent a right anterior sectionectomy. After the operation, bile leakage from the right inferior intrahepatic duct was detected. An endoscopic approach failed because a guidewire could not be passed from the CBD to the right inferior intrahepatic duct. A percutaneous transhepatic approach also failed because the guidewire could not be inserted across the bile leakage site. Radiologic-guided percutaneous transhepatic biliary drainage (PTBD) was performed at the right inferior intrahepatic duct, which reduced the amount of percutaneous bile drainage. However, after PTBD clamping, the amount of PCD increased, and the PTBD catheter

| Table 1. Baseline characteristics of the 20 cases |
|-----------------------------------------------|
| Characteristic | Value |
| Age, yr | 51.4 ± 14.1 |
| Sex, male/female | 18/2 (90/10) |
| Cause of operation | |
| Acute cholecystitis | 13 (65) |
| Hepatocellular carcinoma | 5 (25) |
| Rectosigmoid carcinoma with liver metastasis | 2 (10) |
| Types of surgery | |
| Laparoscopic cholecystectomy | 13 (65) |
| Segmentectomy | 4 (20) |
| Lobectomy | 2 (10) |
| Partial resection | 1 (5) |
| Mode of presentation | |
| Bile drainage through a Jackson-Pratt bag | 15 (75) |
| Abdominal pain | 4 (20) |
| Fever | 1 (5) |
| Site of bile duct injury | |
| Cystic duct stump | 10 (50) |
| Right/left intrahepatic ducts | 3/2 (25) |
| Liver bed | 3 (15) |
| Common hepatic duct | 1 (5) |
| Common bile duct | 1 (5) |
| Types of injury | |
| Leak only | 12 (60) |
| Leak combined with ductal stricture | 3 (15) |
| Leak combined with common bile duct stone | 5 (25) |
| Interval period, day | |
| From surgery to diagnosis of the bile leak | 9.0 ± 6.4 |
| Operation to ERCP | 12.4 ± 8.5 |
| ERCP to stent removal | 77.6 ± 21.7 |

Values are presented as mean ± SD or number (%).
ERCP, endoscopic retrograde cholangiopancreatography.

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could not be removed. The PCD catheter was removed 2 weeks later. In this case, reoperation was judged in the previous hospital to be difficult, with which our surgeon agreed. Therefore, this patient was discharged with PTBD and transferred to another hospital. At the other hospital, the stricture was resolved by positioning the magnet at the stricture site via PTBD and ERCP (Table 2).

DISCUSSION

Postoperative bile leakage is a severe clinical problem that may cause significant morbidity and mortality. Despite improvements in hepatobiliary surgical techniques and postoperative patient care, the incidence of postoperative bile leakage has not decreased significantly [8].

When bile leakage is suspected during the postoperative period, visualization of the biliary tract by ERCP is required to confirm the diagnosis, to locate the exact site of the defect, and to rule out the presence of retained biliary stones.

Bile leakage is more frequent after laparoscopic cholecystectomy than after open cholecystectomy, and the most frequent site of bile leakage is the cystic duct stump [1,3]. Studies of endoscopic treatment for bile leaks of the cystic stump after laparoscopic cholecystectomy have reported promising results. Indeed, in our study endoscopic therapy for bile leaks after laparoscopic cholecystectomy yielded a 100% success rate.

Endoscopic treatment for bile leakage after liver resection shows a lower success rate (75%) than that for bile leakage after laparoscopic cholecystectomy (70% to 100%) [14,16]. In the failed case in our study, peripheral bile leakage occurred because of disconnection between the right posterior intrahepatic duct and common hepatic duct during segmentectomy. In this case, both an endoscopic and percutaneous approach failed. Donatelli et al. [17] proposed the rendezvous technique (passage of a guidewire through the distal CBD and into the subhepatic space by ERCP, and simultaneous snaring via a percutaneous transhepatic cholangioscopy approach allowed for a biliary-duodenal catheter to be placed and the continuity of the biliary tree to be maintained) for this difficult case.

Available endoscopic procedures for bile leakage include EST, endoscopic nasobiliary drainage (ENBD), ERBD with or without EST, or a combination of these procedures. ENBD was not performed in our study because of its inconvenience and risk of accidental tube removal. EST alone is not a good option for treatment of postoperative bile leakage. Marks et al. [18] reported, in a randomized and controlled study using a canine model, that biliary stenting facilitates more rapid resolution of leakage than does EST alone. The theoretical mechanism of resolution of bile leakage is lowering the transpapillary pressure gradient through ampullary orifice and bridging of the leak site. EST alone can lead to equalization of the pressure gradient between the bile duct and duodenum, which decreases flow resistance and diverts bile flow into the duodenum, consequently facilitating leak sealing [3,19]. After EST, placement of a plastic stent at the leak site offers a more predictable pressure gradient-lowering effect and prevents mucosal damage due to bile contents by means of a bridging effect. In addition, stenting can induce bile flow preferentially toward the ampulla [3,20-22]. According to Kaffes et al. [13], endoscopic therapy was carried out in 97 of 100 patients with bile leakage. The endoscopic procedures included stent insertion alone (40 patients), sphincterotomy alone (18 patients), and the combination of both procedures (31 patients). The number of treatment failures was significantly greater in the EST-only group than in the stent-only and combination groups [13]. In our study, the optimum treatment of bile leakage after hepatobiliary surgery was EST plus biliary stenting, which was performed in all cases of postoperative bile leakage. Thirteen cases of bile leakage occurred after laparoscopic cholecystectomy, and all underwent successful endoscopic treatment. One of the seven cases in which bile leakage occurred after liver resection experienced treatment failure, resulting in a failure rate of 14.2% in this study.

Table 2. Complications and failures

| Parameter                                      | No. (%) |
|------------------------------------------------|---------|
| ERBD-related complication                      |         |
| Microperforation during cannulation             | 1 (5)   |
| Failure                                         |         |
| Failure of guide-wire insertion at the bile leak site | 1 (5)   |

ERBD, endoscopic retrograde biliary drainage.
The length, size, and type of plastic stent used varied. Plastic stents of various lengths (5 cm in 4.5%, 7 cm in 13.6%, 9 cm in 9.1%, 10 cm in 41%, 12 cm in 22.7%, and 15 cm in 9.1%), sizes (7 Fr in 86.4% and 10 Fr in 13.6%), and types (straight in 31.5% and double pig tail in 68.5%) were inserted at the sites of bile leakage (Table 3). However, the optimum length, size and type of plastic stent, and the time required for complete resolution have not been determined. A large review of endoscopic intervention for biliary leaks after laparoscopic cholecystectomy by the Midwest Pancreatico Biliary Group concluded that the diameter of the stent does not appear to be important for the treatment of biliary leaks, and 7 Fr stents are a sufficient size for that purpose [15]. A similar conclusion was reached by a large Australian retrospective review of endoscopic intervention in patients with postcholecystectomy bile leakage [13]. In our study, the length, size, and type of plastic stents inserted did not affect the treatment outcomes.

Bile leakage sealing generally takes 1 week after stent placement, and complete closure of the leak site takes 4 to 7 weeks. Therefore, it is usually recommended to remove plastic stents 4 to 6 weeks after placement [2,4,19,22,23]. The average stent maintenance period was 10 to 12 weeks (mean, 77.6 ± 21.7 days) in our study, which is longer than that in previous reports. Nevertheless, no stent malfunction occurred during the maintenance period, and good treatment outcomes were reported.

Based on our results, the overall success rate of ERBD with EST for treatment of bile leakage was 95%. Furthermore, there was no severe endoscopic procedure-induced complication. In conclusion, endoscopic therapy is safe and effective for management of hepatobiliary surgery-induced bile leakage. The endoscopic approach should be considered the first-line modality for the diagnosis and treatment of bile leakage after hepatobiliary surgery.

**KEY MESSAGE**

1. Bile leakage can occur after hepatobiliary surgery and can lead to serious complications.
2. Endoscopic treatment is a safe and effective method for management of hepatobiliary surgery-induced bile leakage

**Conflict of interest**

No potential conflict of interest relevant to this article was reported.

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