

[ORIGINAL ARTICLE]

Can Trainees Safely Perform Endoscopic Treatments for Common Bile Duct Stones? A Single-center Retrospective Study

Tomoya Iida¹, Hiroyuki Kaneto¹, Kohei Wagatsuma¹, Hajime Sasaki¹, Yumiko Naganawa¹, Suguru Nakagaki¹, Shuji Satoh¹, Haruo Shimizu¹ and Hiroshi Nakase²

Abstract:

Objective There are no reports on whether or not trainees can safely carry out endoscopic procedures for the removal of common bile duct (CBD) stones. The aim of this study was to investigate the efficacy and safety of endoscopic treatments for CBD stones by trainees.

Methods Endoscopic retrograde cholangiopancreatography (ERCP) was performed in 1,016 consecutive patients at our institution during the 6-year study period. The endoscopically treated patients with CBD stones were included in this study. Physicians who had experienced \geq 300 ERCP procedures were defined as experts, while those who had experienced <300 procedures were defined as trainees. The trainees were replaced by an expert when they could not achieve the established criteria. Patients were divided into the following three groups to retrospectively examine the patients' backgrounds, details of endoscopic treatments, and intra-/post-operative complications: Group A, completed by trainees under supervision of an expert; B, treated by an expert who switched in for a trainee in the middle of the procedure; and C, completed by an expert.

Results A total of 325 patients with CBD stones underwent endoscopic treatments. The number included in Groups A, B, and C was 176, 102, and 47, respectively. The bile duct catheter insertion successes rates for Groups A, B, and C were 99.0%, 97.1%, and 100% (p=0.09), and the complete stone removal rates were 94.2%, 94.8%, and 100%, respectively (p=0.07), showing no significant difference among the three groups. Furthermore, the frequency of intra-/post-operative complications was not significantly different among the three groups (p=0.48, p=0.12, respectively).

Conclusion This study showed that trainees could safely perform endoscopic procedures in accordance with our facility's criteria during ERCP.

Key words: endoscopic retrograde cholangiopancreatography (ERCP), endoscopy, common bile duct, stone, trainee

(Intern Med 57: 923-928, 2018) (DOI: 10.2169/internalmedicine.9737-17)

Introduction

Common bile duct (CBD) stones are a frequently encountered disease in clinical practice and constitute a gastrointestinal emergency disease, as they result in cholangitis and sepsis. The endoscopic management of CBD stones has been regarded as less invasive than surgery (1-3). Endoscopic treatments are therefore considered the first choice for the management of CBD stones.

In recent years, the Tokyo Guideline derived from international meetings in 2007 (4) and updated in 2013 (5) was published for the diagnosis, classification, and treatment of acute cholangitis. Although treatment is defined by severity in the Tokyo Guideline, endoscopic treatment has an important role in all severities.

¹Department of Gastroenterology, Muroran City General Hospital, Japan and ²Department of Gastroenterology and Hepatology, Sapporo Medical University School of Medicine, Japan

Received: June 28, 2017; Accepted: August 7, 2017; Advance Publication by J-STAGE: February 9, 2018 Correspondence to Dr. Hiroshi Nakase, hiropynakase@gmail.com

Endoscopic retrograde cholangiopancreatography (ERCP) is a relatively difficult procedure, and whether or not trainees can perform it safely remains unclear (6-8). Furthermore, no reports have described whether or not trainees can safely carry out procedures for the removal of CBD stones. The aim of this study was to investigate the efficacy and safety of endoscopic treatments for CBD stones performed by trainees.

Materials and Methods

Patients

ERCP was performed in 1,016 consecutive patients at our institution during a 6-year study period (January 2009 to December 2014). The endoscopically treated patients were included in this study. The study design was approved by the local ethics committee, and all patients signed a standard informed consent form before undergoing the endoscopic procedure.

Endoscopic procedure

First, physicians who had experienced \geq 300 ERCP procedures were defined as experts, while those who had experienced <300 procedures were defined as trainees. ERCP was performed by an expert operator (H.K.) and three trainees (T.I., H.S., and Y. N.) under the supervision of the expert.

Physicians who had been mainly in charge of patients started the endoscopic procedure regardless of difficulty level. Trainees started the procedure and switched with the expert physician when they were unable to achieve viewing from the front of the papilla within 5 minutes, cannulation within 15 minutes, and completion within 60 minutes from 30 minutes after starting the procedure.

Pentazocine and midazolam (MDZ) were administered intravenously for conscious sedation under monitoring of peripheral oxygen saturation and blood pressure throughout the procedure. The initial dosages of pantazocine and MDZ were 15 mg and 1.25-2.5 mg, respectively, and the dose was adjusted according to the condition of the patient. Oxygen supplementation through nasal cannulae was used as necessary. Vital signs, dosages of sedatives, and the responses of patients were recorded in detail by nurses during the procedure.

Standard duodenoscopes (JF-260V and TJF-240; Olympus Medical Systems, Tokyo, Japan) were used for the procedures. Furthermore, we used 2T-2Q260M and CF-Q240AI (Olympus Medical Systems) for patients who had undergone gastric surgery.

Guidewire cannulation was used for bile duct catheter insertion (9). When a biliary approach failed using a standard method, a precut technique using a needle knife (Boston Scientific, Tokyo, Japan) was attempted as a rescue technique. Endoscopic sphincterotomy (EST) (2), endoscopic papillary balloon dilation (EPBD) (10), or endoscopic papillary large balloon dilation (EPLBD) (11) was performed to facilitate stone removal in most cases. While EST (medium incision or larger) was carried out for the first papillotomy, EPBD was selected depending on the use of anti-thrombotic drugs and the condition of the periampullary diverticulum. EPLBD was selected for patients with 3 or more stones and with a sufficiently dilated bile duct from the lower bile duct to the hilar bile duct, even if the stones were 12 mm or longer or about 10 mm in diameter.

For stone removal, standard techniques (basket, balloon catheter, mechanical lithotripsy) were applied. A mechanical lithotriptor was used to crush stones too large to retrieve intact. Extracorporeal shock wave lithotripsy (ESWL) was used for large stones that could not be crushed by mechanical lithotripsy. Essentially, complete duct clearance was attempted at each procedure. The need for repeat endoscopy was determined on discussion with the patient considering the stability of his/her general condition during the previous procedure and the likelihood of complete duct clearance. When complete stone extraction was not achieved, a 7.0-Fr straight or double-pigtail catheter (Flexima[™] Biliary Stent System, Boston Scientific, Marlborough, USA) was positioned for bile duct drainage. Stent exchange was performed only when acute cholangitis recurred.

After the procedure, we administered 0.2-0.5 mg flumazenil. As a preventive measure against pancreatitis after ERCP, a sufficient amount of replacement fluid was given in addition to administration of protease inhibitors and ulinastatin in all cases. Furthermore, immediately after ERCP procedures, patients received 25-50 mg diclofenac suppositories depending on the judgment of the operator.

Examination items

The proportion who underwent an endoscopic procedure, number of patients by group, patient background [age, sex, Eastern Cooperative Oncology Group performance status scale (PS), presence or absence of concomitant diseases/ cholecystectomy/periampullary diverticulum/history of previous treatments/gastrectomy/anti-thrombotic drugs], details of endoscopic therapy (complete removal rate of CBD stones, procedure time, procedure number, dose of MDZ, long diameter of CBD, short diameter and number of stones, papillary treatment, use of a pancreatic duct guidewire, and placement of a pancreatic duct stent), and complications were reviewed based on medical charts and films. We investigated the complications that occurred during and after ERCP. Complications during the procedure were cardiorespiratory suppression, and those after procedure were pancreatitis, cholangitis, and bleeding. Cardiorespiratory suppression was defined as peripheral oxygen saturation <90% and/or systolic blood pressure <90 mmHg observed at any time during the procedure. Post-ERCP pancreatitis (PEP), cholangitis, bleeding and their severities were determined according to the 1991 consensus guidelines by Cotton et al. (12)

The above-mentioned items were examined by dividing patients into the following three groups: Group A, treated by



Figure. Study protocol. ERCP was performed in 1,016 consecutive patients at our institution during the 6-year study period. A total of 325 patients with CBD stones underwent ERCP. Physicians who had experienced \geq 300 ERCP procedures were defined as experts, while those who had experienced <300 procedures were defined as trainees. Patients were divided into the following three groups: Group A, treated by trainees (completed by trainees from insertion to removal of endoscopy under supervision of an expert); Group B, treated by an expert who switched in for a trainee in the middle of the procedure (operator switch-in based on the abovementioned criteria); and Group C, treated by an expert (completed by an expert from insertion to removal of endoscopy). The number of cases included in Group A, Group B, and Group C was 176, 102, and 47, respectively.

trainees (completed by trainees from insertion to removal of endoscopy under supervision of an expert); Group B, treated by an expert who switched in for a trainee in the middle of the procedure (operator switch-in based on the abovementioned criteria); and Group C, treated by an expert (completed by an expert from insertion to removal of endoscopy) (Figure).

Data analyses

Data were recorded and analyzed with EZR (13) version 1.33. Results were reported as the mean and standard deviation for variables with a normal distribution, and as the median, minimum, and maximum for variables with a nonnormal distribution. A univariate analysis was conducted with the chi-squared test for nominal variables, with nonparametric tests (Mann-Whitney U test, Kruskall-Wallis test) for variables with a non-normal distribution, and with Student's *t*-test for variables with a normal distribution. Two-sided hypothesis testing was used, with a p value of less than 0.05 considered statistically significant.

Results

During the study period, 325 patients with CBD stones underwent ERCP (32.0% of 1,016 ERCP procedures during the same period). The overall bile duct catheter insertion success rate was 98.8%, and the complete stone removal rate was 98.5%. The number of cases included in Group A, Group B, and Group C were 176, 102, and 47, respectively, the bile duct catheter insertion success rates were 99.0%, 97.1%, and 100%, respectively (p=0.09), and the complete stone removal rates were 94.2%, 94.8%, and 100%, respectively (p=0.07), showing no significant difference among the three groups.

No significant difference was observed in age, sex, or PS among three groups. Furthermore, no significant difference was found in overall concomitant diseases, cardiovascular disease, cerebrovascular, metabolic diseases, dementia, hemodialysis, periampullary diverticulum, or the use of antithrombotic drugs, anti-platelet drugs, or anti-coagulant drugs (Table 1). No significant differences in the CBD diameter, the stone diameter, the number of stones, the procedure time, or the procedure details (EST, EPBD, EPLBD, and precut) were observed among the three groups (Table 2). The frequency of intra-operative complications (cardiorespiratory suppression) and post-operative complications (pancreatitis, cholangitis, bleeding) was not significantly different among the three groups. With regard to post-operative complications, there were no significant differences in the severity of either pancreatitis or cholangitis (Table 3).

In Group B, patients more frequently had received the procedure as a first-line therapy (A:B:C= 63.0%:79.4%: 61.7%; p=0.01), had a history of papillary procedure (A:B:C= 36.9%:20.6%:38.3%; p=0.0028), had undergone gastrectomy (2.3%:13.7%:6.4%; p=0.0009), and had a history of malignant disease than the other two groups (14.2%:27.5%: 21.3%; p=0.025). In addition, the dose of MDZ [5.8 mg (1.25-20 mg):6.5 mg (1.25-17.5):5.3 mg (1.25-12.5 mg)] was larger (p=0.048) and the procedure time longer [36 minutes (10-90 minutes):46 minutes (19-118 minutes):30 minutes (15-108 minutes)] (p<0.0001) in this group than in the other two groups (Table 1, 2).

Discussion

This retrospective study aimed to evaluate whether or not

Table 1. Patients Backgrounds.

	Group A (n=176)		Group B (n=102)		Group C (n=47)		p value
Age (years) (range)	75 (29-96)		76 (34-96)		77 (45-89)		0.98
Sex	M: F=98: 78		M: F=56: 46		M: F=25: 22		0.95
PS median (range)	1 (0-3)		1 (0-4)		1 (0-4)		0.57
Presence/ Absence	+	-	+	-	+	-	
Concomitant disease (%)	88.1	11.9	93.1	6.9	87.2	12.8	0.35
History of malignant diseases (%)	14.2	85.8	27.5	72.5	21.3	78.7	0.025
Cardiovascular diseases (%)	56.8	43.2	62.7	27.3	63.8	36.2	0.51
Cerebrovascular diseases (%)	27.8	72.2	21.6	78.4	29.8	70.2	0.43
Metabolic diseases (%)	33.5	66.5	24.5	75.5	27.7	72.3	0.27
Dementia (%)	4.5	95.5	10.8	89.2	6.4	93.6	0.14
Hemodialysis (%)	5.7	94.3	2.9	97.1	2.1	97.9	0.50
Other diseases (%)	26.1	73.9	22.5	77.5	14.9	85.1	0.26
Cholecystectomy (%)	29.5	70.5	19.6	80.4	46.8	53.2	0.0029
Periampullary diverticulum (%)	44.9	55.1	41.1	58.8	48.9	51.1	0.66
Previous procedures for the papilla (%)	36.9	63.1	20.6	79.4	38.3	61.7	0.0028
Gastrectomy (%)	2.3	97.7	13.7	86.3	6.4	93.6	0.0009
Anti-thrombotic drugs (%)	33.5	66.5	38.2	61.8	34.0	66.0	0.72
Anti-platelet drugs (%)	27.8	73.2	28.4	71.6	31.9	68.1	0.86
Thienopyridine series (%)	5.6	94.4	3.9	96.1	2.1	97.9	0.62
Low-dose aspirin (%)	21.6	78.4	22.5	77.5	25.5	74.5	0.85
Anti-coagulant drugs (%)	8.0	92.0	11.8	88.2	2.1	97.9	0.14

PS: Eastern Cooperative Oncology Group performance status scale

Table 2. Contents of Endoscopic Treatments.

	Group A (n=176)		Group I (n=102)		Group ((n=47)	2	p value
Complete removal of CBD stones rate (%)	94.2		94.8		100		0.07
Bile duct catheter insertion successes rate (%)	99		97.1		100		0.08
MDZ (mean) (range)	5.8mg (1.25-20)		6.5mg (1.25-17.5)		5.3mg (1.25-12.5)		0.048
Procedure time (mean) (range)	36 min (10-90)		46min (19-118)		30min (15-108)		< 0.0001
Long diameter of CBD (mean) (range)	9.4mm (4.6-20.1)		9.7mm (4.8-14.8)		9.8mm (5.2-32.0)		0.94
Number of CBD stones (median) (range)	1 (0-12)		2 (0-10)		2 (0-12)		0.48
Short diameter of CBD stones (mean) (range)	7.0mm (2.5-19.5)		7.5mm (2.8-20.5)		6.8mm (3.4-34.0)		0.57
Presence/ Absence	+	-	+	-	+	-	
Only ENBD tube inserted (%)	5.1	94.9	4.9	95.1	0	100	0.36
Procedures for the papillia (%)	70.5	29.5	84.3	15.7	68.1	31.9	0.02
EST (%)	56.8	43.2	68.6	31.4	57.4	42.6	0.14
EPBD (%)	11.9	88.1	14.7	85.3	10.6	89.4	0.78
EPLBD (%)	6.2	93.8	11.8	88.2	0	100	0.10
Pre-cut (%)	0	100	2.9	97.1	0	100	0.053

CBD: common bile duct, MDZ: midazolam, ESWL: extracorporeal shock wave lithotripsy, ENBD: endoscopic nasobilliary drainage, EST: endoscopic sphincterotomy, EPBD: endoscopic papillary balloon dilation, EPLBD: endoscopic papillary large balloon dilation

trainees could safely perform endoscopic treatment for CBD stones under experts' instruction. Our data showed that trainees could safely perform these procedures according to our facility's criteria and rules during ERCP because of the lack of a significant difference in the success rate of the bile duct catheter insertion, the complete stone removal rate, and the intra-/post-operative complication among Groups A (treated by trainees), B (experts switched in for trainees in the middle of the procedure), and C (treated by experts).

form ERCP is an issue, as complications with ERCP are more numerous and severe than with other endoscopic procedures. There are several reports concerning the safety of ERCP performed by trainees. Voiosu et al. (7) reported that the ERCP technical success rate increased with trainee experience, reflecting the learning curve of individual operators in their study. In addition, the complication rates were similar despite different levels of operative experience. In contrast, Ekkelenkamp et al. (8) demonstrated a relationship between the involvement of trainees during ERCP and failure

Educating young gastroenterologists on how to safely per-

	Group A (n=176)	Group B (n=102)	Group C (n=47)	p value
Complications	7.4	9.8	12.8	0.48
during ERCPs (%)				
Complications	13.1	20.6	8.5	0.12
post ERCPs (%)				
Pancreatitis (All) (%)	11.4	14.7	8.5	0.57
Pancreatitis (Mild/Medium/Severe) (%)	9.7/1.7/0	8.8/3.9/2.0	6.4/0/2.1	0.28
Cholangitis (All) (%)	1.7	3.9	0	0.30
Cholangitis (Mild/Medium/Severe) (%)	1.7/0/0	2.0/2.0/0	0/0/0	0.18
Bleeding (All) (%)	0.6	3.9	0	0.06
Bleeding (Mild/Medium/Severe) (%)	0.6/0/0	2.0/2.0/0	0/0/0	0.20

Table 3.Adverse Events during or Post ERCP.

ERCP: Endoscopic retrograde cholangiopancreatography

of the endoscopic procedure as an independent risk factor in their prospective study.

Nevertheless, we must consider that the safety and success of ERCP performed by trainees might vary depending on patients' general condition and disease status, such as the malignancy status. We therefore focused on the safety of endoscopic treatment for CBD stones performed by trainees and evaluated the outcomes by dividing patients into Groups A (treated by trainees), B (experts switched in for trainees in the middle of the procedure), and C (treated by an expert). No previous study has conducted an investigation from this viewpoint.

A recent systematic review (6) suggested that ERCP trainees might require experience with 70-400 procedures to achieve a certain level of success in bile duct catheter insertion (14-22). In reference to this review, in our facility, we defined physicians who had performed \geq 300 ERCP procedures as experts.

In addition, the recent European Society of Gastrointestinal Endoscopy guideline (23) suggests that the supervisor should take over the procedure from the trainee when the papilla is deemed difficult to cannulate. Despite the lack of a precise definition for "difficult biliary cannulation", a prospective multicenter study by Testoni et al. (24) showed a linear progression between ≤3 and 4-10 attempts and between 4-10 and >10 attempts. According to a previous metaanalysis (25), cannulation attempts of >10 minutes' duration represented an independent risk factor with an odds ratio (OR) of 1.76 [95% confidence interval (CI): 1.13-2.74], and the pooled incidence of PEP increased from 3.8% to 10.8% compared with cannulation attempts of ≤10 minutes' duration. A recent report also indicated that the appropriate cannulation time for trainees is 10 minutes (26). Based on these reports, the rule for switching from trainees to experts during ERCP in our facility was decided as follows: (1) viewing from the front of the papilla should be completed within 5 minutes, (2) cannulation should be completed within 15 minutes, and (3) the procedure should be completed within 60 minutes.

The present data suggested that trainees could safely per-

form ERCP for CBD stones under instruction by experts with our facility's definition and switching rule. In clinical practice, many physicians in Group B were concerned about the clinical outcome (procedure time, complications associated with ERCP). Based on the patients' background, Group B seemed to include many endoscopically difficult cases. It was therefore natural that Group B included a higher ratio of pancreatic duct pancreatic duct guidewire placement, higher dosage of MDZ, and longer procedure time than Groups A and C. However, no significant difference in the intra-/post-operative complication rate was observed among all groups, suggesting that our facility's definition and switching rule are reliable for trainees performing ERCP for CBD stones.

Several limitations associated with the present study warrant mention. First, this was a single-center, retrospective study with a small number of enrolled patients. Second, there might be bias in patients' background data, such as the significantly higher rate of patients with a history of cholecystectomy in Group C than in the other two groups. Third, we were unable to evaluate the cardiac function for all groups in detail, although there was no marked difference in the ratio of cardiac diseases among the groups.

In conclusion, this study demonstrated that trainees could safely carry out endoscopic treatment for CBD stones under our facility's definitional switching rule. However, a multicenter prospective study will be required to confirm the validation of our educational system.

The authors state that they have no Conflict of Interest (COI).

References

- 1. Siegel JH, Kasmin FE. Biliary tract diseases in the elderly: management and outcomes. Gut 41: 433-435, 1997.
- Kawai K, Akasaka Y, Murakami K, Tada M, Koli Y. Endoscopic sphincterotomy of the ampulla of Vater. Gastrointest Endosc 20: 148-151, 1974.
- Dasari BV, Tan CJ, Gurusamy KS, et al. Surgical versus endoscopic treatment of bile duct stones. Cochrane Database Syst Rev 9: CD003327, 2013.

- Takada T, Kawarada Y, Nimura Y, et al. Tokyo Guidelines for the management of acute cholangitis and cholecystitis. J Hepatobiliary Pancreat Surg 14: 1-10, 2007.
- Takada T, Strasberg SM, Solomkin JS, et al. TG13: Updated Tokyo Guidelines for the management of acute cholangitis and cholecystitis. J Hepatobiliary Pancreat Surg 20: 1-7, 2013.
- Shahidi N, Ou G, Telford J, Enns R. When trainees reach competency in performing ERCP: a systematic review. Gastrointest Endosc 81: 1337-1342, 2015.
- Voiosu T, Bengus A, Voiosu A, et al. Trainee caseload correlates with ERCP success but not procedure-related complications: results from a prospective study (the QUASIE cohort). Endosc Int Open 4: E409-E414, 2016.
- **8.** Ekkelenkamp VE, de Man RA, Ter Borg F, et al. Prospective evaluation of ERCP performance: results of a nationwide quality registry. Endoscopy **47**: 503-507, 2015.
- Lella F, Bagnolo F, Colombo E, et al. A simple way of avoiding post-ERCP pancreatitis. Gastrointest Endosc 59: 830-834, 2004.
- 10. Staritz M, Ewe K, Meyer zum Büschenfelde KH. Endoscopic papillary dilation (EPD) for the treatment of common bile duct stones and papillary stenosis. Endoscopy 15 (Suppl 1): 197-198, 1983.
- Ersoz G, Tekesin O, Ozutemiz AO, Gunsar F. Biliary sphincterotomy plus dilation with a large balloon for bile duct stones that are difficult to extract. Gastrointest Endosc 57: 156-159, 2003.
- 12. Cotton PB, Lehman G, Vennes J, et al. Endoscopic sphincterotomy complications and their management: an attempt at consensus. Gastrointest Endosc 37: 383-393, 1991.
- Kanda Y. Investigation of the freely available easy-to-use software 'EZR' for medical statistics. Bone Marrow Transplant 48: 452-458, 2013.
- 14. Jowell PS, Baillie J, Branch S, et al. Quantitative assessment of procedural competence: a prospective study of training in endoscopic retrograde cholangiopancreatography. Ann Intern Med 125: 983-989, 1996.
- Watkins JL, Etzkom KP, Wiley TE, DeGuzman L, Harig JM. Assessment of technical competence during ERCP training. Gastrointest Endosc 44: 411-415, 1996.
- Schlup MM, William SM, Barbezat GO. ERCP: a review of technical competency and workload in a small unit. Gastrointest Endosc 46: 48-52, 1997.

- Biau DJ, William SM, Schlup MM, Nizard RS, Porcher R. Quantitative and individualized assessment of the learning curve using LC-CUSUM. Br J Surg 95: 925-929, 2008.
- Kowalski T, Kanchana T, Pungpapong S. Perceptions of gastroenterology fellows regarding ERCP competency and training. Gastrointest Endosc 58: 345-349, 2003.
- Vitale GC, Zavaleta CM, Vitale DS, Binford JC, Tran TC, Larson GM. Training surgeons in endoscopic retrograde cholangiopancreatography. Surg Endosc 20: 149-152, 2006.
- 20. Verma D, Gostout CJ, Petersen BT, Levy MJ, Baron TH, Adler DG. Establishing a true assessment of endoscopic competence in ERCP during training and beyond: a single-operator learning curve for deep biliary cannulation in patients with native papillary anatomy. Gastrointest Endosc 65: 394-400, 2007.
- Waller HM, Connor SJ. Cumulative sum (Cusum) analysis provides an objective measure of competency during training in endoscopic retrograde cholangio-pancreatography (ERCP). HPB (Oxford) 11: 565-569, 2009.
- 22. Ekkelenkamp VE, Koch AD, Rauws EA, Borsboom GJ, de Man RA, Kuipers EJ. Competence development in ERCP: the learning curve of novice trainees. Endoscopy 46: 949-955, 2014.
- 23. Testoni PA, Mariani A, Aabakken L, et al. Papillary cannulation and sphincterotomy techniques at ERCP: European Society of Gastrointestinal Endoscopy (ESGE) Clinical Guideline. Endoscopy 48: 657-683, 2016.
- 24. Testoni PA, Mariani A, Giussani A, et al. Risk factors for post-ERCP pancreatitis in high- and low-volume centers and among expert and non-expert operators: a prospective multicenter study. Am J Gastroenterol 105: 1753-1761, 2010.
- 25. Masci E, Mariani A, Curioni S, Testoni PA. Risk factors for pancreatitis following endoscopic retrograde cholangiopancreatography: a meta-analysis. Endoscopy 35: 830-834, 2003.
- 26. Pan Y, Zhao L, Leung J, et al. Appropriate time for selective biliary cannulation by trainees during ERCP-a randomized trial. Endoscopy 47: 688-695, 2015.

The Internal Medicine is an Open Access article distributed under the Creative Commons Attribution-NonCommercial-NoDerivatives 4.0 International License. To view the details of this license, please visit (https://creativecommons.org/licenses/by-nc-nd/4.0/).

© 2018 The Japanese Society of Internal Medicine Intern Med 57: 923-928, 2018