

Response:

We appreciate Ding et al's¹ interest and comments related to our video case report entitled "Endoclip papilloplasty for a patulous and incompetent biliary papilla: a therapeutic misadventure."² In their letter, they claimed that clipping is likely to increase the likelihood of adverse events and that biliary-enteric anastomosis has been shown to be safe.

It is well known that endoscopic hemostasis by clipping is safe and effective to manage refractory postsphincterotomy bleeding and tear. In addition, these clips generally fall off spontaneously in days or weeks. In our experience, endoclips fell off within 3 weeks in 70% of patients who had undergone endoclip papilloplasty (30 patients, unpublished data). By contrast, rates of late adverse events after biliary-enteric anastomosis, such as biliary or anastomotic strictures, were reported to be 10% to 30%.^{3,4} These patients are susceptible to recurrent choledocholithiasis and cholangitis. Therefore, we believe that it is incorrect to claim there is no need to preserve the sphincter of Oddi (SO) function.

Biliary sludge contains bacteria and fungi, microbial by-products, proteins, dietary fibers, crystals of fatty acid calcium salts, and amorphous calcium bilirubinate.⁵⁻⁷ The SO prevents the reflux of duodenal contents into the bile duct, the pancreatic duct, or both, under physiologic conditions.⁸ This natural antireflux function disappears after sphincterotomy.⁹ Permanent destruction of the SO after endoscopic sphincterotomy can lead to late adverse events such as recurrent choledocholithiasis, ascending cholangitis, and acute cholecystitis.¹⁰ The risks of these late adverse events are related to long-standing duodenobiliary reflux with bacterial contamination.¹¹ Although risk factors related to common bile duct stone recurrence are multiple, including the differences in stone ingredients,¹² biliary microbiota,^{13,14} genetic factors,^{15,16} juxtapapillary diverticulum,¹⁷ and biliary duct dilation,¹⁸ endoscopic sphincterotomy, endoscopic papillary large balloon dilation, and biliary microbiota are clearly related to stone recurrence.

Although a scarred or repaired SO may lack certain neuronally controlled muscle fibers, the control of the SO is complex, and it is under hormonal, extrinsic neural, intrinsic neural, and myogenic control or influence.¹⁹ Our porcine experiments (unpublished data) demonstrated that healing of the SO after endoclip papilloplasty was different from that in the control group. We also demonstrated that the basic pressure of the SO in the treatment group was significantly higher than that in the control group.²⁰

We concur that endoclip papilloplasty is not suitable for a papilla located deep within the diverticulum, a small papilla, and a papilla with a short intramural segment. These papillas are not suitable for extensive sphincterotomy anyway. We also agree that long-term follow-up and

large-scale randomized controlled trials are needed in this direction.

Yong-hui Huang, MD
Peking University Third Hospital
Beijing, China

REFERENCES

- Ding H, Rajman I, Kalloo AN, et al. Endoclip papilloplasty for a patulous and incompetent biliary papilla: a therapeutic misadventure. VideoGIE 2019;4:493.
- Fan X, Li X, Chang H, et al. Endoclip papilloplasty for a patulous and incompetent biliary papilla. VideoGIE 2019;4:331-3.
- Perera MT, Silva MA, Hegab B, et al. Specialist early and immediate repair of post-laparoscopic cholecystectomy bile duct injuries is associated with an improved long-term outcome. Ann Surg 2011;253:553-60.
- Walsh RM, Henderson JM, Vogt DP, et al. Long-term outcome of biliary reconstruction for bile duct injuries from laparoscopic cholecystectomies. Surgery 2007;142:450.
- Groen AK, Out T, Huibregts K, et al. Characterization of the content of occluded biliary endoprostheses. Endoscopy 1987;19:57-9.
- Speer AG, Cotton PB, Rode J, et al. Biliary stent blockage with bacterial biofilm: a light and electron microscopy study. Ann Intern Med 1988;108:546-53.
- Leung JW, Ling TK, Kung JL, et al. The role of bacteria in the blockage of biliary stents. Gastrointest Endosc 1988;34:19-22.
- Liu YF, Saccone GT, Thune A, et al. Sphincter of Oddi regulates flow by acting as a variable resistor to flow. Am J Physiol 1992;263:G683-9.
- Sung JY, Leung JW, Shaffer EA, et al. Ascending infection of the biliary tract after surgical sphincterotomy and biliary stenting. J Gastroenterol Hepatol 1992;7:240-5.
- Tocchi A, Mazzoni G, Liotta G, et al. Late development of bile duct cancer in patients who had biliary-enteric drainage for benign disease: a follow-up study of more than 1,000 patients. Ann Surg 2001;234:210-4.
- van Berkel AM, van Marie J, van Veen H, et al. A scanning electron microscopic study of biliary stent materials. Gastrointest Endosc 2000;51:19-22.
- Grühnäge F, Lammert F. Pathogenesis of gallstones: a genetic perspective. Best Pract Res Clin Gastroenterol 2006;20:997-1015.
- Kose SH, Grice K, Orsi WD, et al. Metagenomics of pigmented and cholesterol gallstones: the putative role of bacteria. Sci Rep 2018;8:11218.
- Wu T, Zhang Z, Liu B, et al. Gut microbiota dysbiosis and bacterial community assembly associated with cholesterol gallstones in large-scale study. BMC Genomics 2013;14:669.
- Rosmorduc O, Hermelin B, Boelle PY, et al. ABCB4 gene mutation-associated cholelithiasis in adults. Gastroenterology 2003;125:452-9.
- Oude Elferink RP, Beuers U. Targeting the ABCB4 gene to control cholesterol homeostasis. Exp Opin Ther Targets 2011;15:1173-82.
- Ueno N, Ozawa Y, Aizawa T. Prognostic factors for recurrence of bile duct stones after endoscopic treatment by sphincter dilation. Gastrointest Endosc 2003;58:336-40.
- Pereira-Lima JC, Jakobs R, Winter UH, et al. Long-term results (7 to 10 years) of endoscopic papillotomy for choledocholithiasis: multivariate analysis of prognostic factors for the recurrence of biliary symptoms. Gastrointest Endosc 1998;48:457-64.
- Tanaka M. Function and dysfunction of the sphincter of Oddi. Dig Surg 2010;27:94-9.
- Huang YH, Wang K, Zhang HJ, et al. The sphincter-preserving effect of duodenal papilla occlusion by SureClip from MicroTech [Abstract]. J Gastroenterol Hepatol 2018;33:118.

<https://doi.org/10.1016/j.vgie.2019.08.010>