

Submucosal Tunneling Endoscopic Resection for Submucosal Tumors of Less Than 35 mm in the Upper and Lower Gastrointestinal Tract: Systematic Review and Meta-Analysis.

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Objectives: We conducted a systematic review and meta-analysis to synthesize the current evidence on the efficacy and safety of submucosal tunneling endoscopic resection (STER) for submucosal tumors (SMTs) of the gastrointestinal (GI) tract (3,4).

Methods: This study adhered to PRISMA guidelines and included observational studies on patients with upper or lower GI tract SMTs who underwent STER (5, 6). Data were extracted from these studies and analyzed using Open Meta[Analyst] and Jamovi software, with outcomes including resection rates, recurrence, and complications. Quality assessment and risk of bias were evaluated using the Newcastle-Ottawa Scale and NIH Quality Assessment Tool, and publication bias was evaluated via funnel plots and regression tests.

Results: This meta-analysis of 27 studies indicated high complete and En-bloc resection rates of 96.1% (95% CI: 94.1% to 98.0%) and 90.3% (95% CI: 85.6% to 94.9%), respectively (1,2,3,7–30). The studies exhibited a low recurrence rate of 2.0% (95% CI: 1.2% to 3.4%) and moderate incidence of complications such as subcutaneous emphysema and pneumomediastinum (10.0%, 95% CI: 5.5% to 17.6%) and pneumothorax (6.6%, 95% CI: 3.9% to 11.0%). Other adverse events like perforation and bleeding were rare, occurring at rates of 2.3% (95% CI: 1.1% to 4.7%) and 2.5% (95% CI: 1.2% to 5.2%), respectively. Furthermore, the average operative time was 59.28 min (95% CI: 53.83 to 64.73), with an average hospital stay of 4.77 days (95% CI: 3.91 to 5.63).

Conclusion: STER demonstrates high resection rates and low recurrence in the treatment of gastrointestinal SMTs. While complications like subcutaneous emphysema, pneumomediastinum, and pneumothorax do occur, they remain low in incidence. Rarer complications, such as perforation and bleeding, underscore the overall safety of the procedure.

Keywords: Submucosal Tunneling Endoscopic Resection; Submucosal Tumors; GIT; Meta-analysis

References

1. Du C, Chai N-L, Ling-Hu EQ, Li ZJ, Li LS, Zou JL, Jiang L, Lu ZS, Meng JY, Tang P. *Submucosal tunneling endoscopic resection: An effective and safe therapy for upper gastrointestinal submucosal tumors originating from the muscularis propria layer*. World J Gastroenterol, 2019. **25**:245–257. doi:10.3748/wjg.v25.i2.245
2. Xu MD, Cai MY, Zhou PH, Qin XY, Zhong YS, Chen WF, Hu JW, Zhang YQ, Ma LL, Qin WZ, Yao LQ. *Submucosal tunneling endoscopic resection: a new technique for treating upper GI submucosal tumors originating from the muscularis propria layer (with videos)*. Gastrointest Endosc, 2012. **75**:195–199.
3. Tu S, Huang S, Li G, Tang X, Qing H, Gao Q, Fu J, Du G, Gong W. *Submucosal Tunnel Endoscopic Resection for Esophageal Submucosal Tumors: A Multicenter Study*. Gastroenterol Res Pract, 2018. **2018**:2149564. doi:10.1155/2018/2149564

4. Dellatore P, Bhagat V, Kahaleh M. *Endoscopic full thickness resection versus submucosal tunneling endoscopic resection for removal of submucosal tumors: a review article*. Transl Gastroenterol Hepatol, 2019. **4**:45. doi:10.21037/tgh.2019.05.03
5. Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, Shamseer L, Tetzlaff JM, Akl EA, Brennan SE, Chou R, Glanville J, Grimshaw JM, Hróbjartsson A, Lalu MM, Li T, Loder EW, Mayo-Wilson E, McDonald S, McGuinness LA, Stewart LA, Thomas J, Tricco AC, Welch VA, Whiting P, Moher D. *The PRISMA 2020 statement: an updated guideline for reporting systematic reviews*. BMJ, 2021. **372**:n71. doi:10.1136/BMJ.N71
6. Chandler J, Higgins JP, Deeks JJ, Davenport C, Clarke MJ. *Handbook for Systematic Reviews of Interventions*. Cochrane Handb Syst Rev Interv Handb Syst Rev Interv Version 520, 2017. 1–11.
7. Wang L, Ren W, Zhang Z, Yu J, Li Y, Song Y. *Retrospective study of endoscopic submucosal tunnel dissection (ESTD) for surgical resection of esophageal leiomyoma*. Surg Endosc, 2013. **27**:4259–4266. doi:10.1007/s00464-013-3035-z
8. Wang H, Tan Y, Zhou Y, Wang Y, Li C, Zhou J, Duan T, Zhang J, Liu D. *Submucosal tunneling endoscopic resection for upper gastrointestinal submucosal tumors originating from the muscularis propria layer*. Eur J Gastroenterol Hepatol, 2015. **27**:776–780. doi:10.1097/MEG.0000000000000394
9. Wang GX, Yu G, Xiang YL, Miu Y De, Wang HG, Xu MD. *Submucosal tunneling endoscopic resection for large symptomatic submucosal tumors of the esophagus: A clinical analysis of 24 cases*. Turkish J Gastroenterol, 2020. **31**:42–48. doi:10.5152/tjg.2020.19062
10. Wang YP, Xu H, Shen JX, Liu WM, Chu Y, Duan BS, Lian JJ, Zhang HB, Zhang L, Xu MD, Cao J. *Predictors of difficult endoscopic resection of submucosal tumors originating from the muscularis propria layer at the esophagogastric junction*. World J Gastrointest Surg, 2022. **14**: 918–929. doi:10.4240/wjgs.v14.i9.918
11. Hu JW, Zhang C, Chen T, Zhou PH, Zhong YS, Zhang YQ, Chen WF, Li QL, Yao LQ, Xu MD. *Submucosal tunneling endoscopic resection for the treatment of rectal submucosal tumors originating from the muscularis propria layer*. J Cancer Res Ther, 2014. **10**:C281–C286. doi:10.4103/0973-1482.151533
12. Ye LP, Zhang Y, Mao XL, Zhu LH, Zhou X, Chen JY. *Submucosal tunneling endoscopic resection for small upper gastrointestinal subepithelial tumors originating from the muscularis propria layer*. Surg Endosc, 2014. **28**: 524–530. doi:10.1007/s00464-013-3197-8
13. Zhang C, Hu JW, Chen T, Zhou PH, Zhong YS, Zhang YQ, Chen WF, Li QL, Yao LQ, Xu MD. *Submucosal tunneling endoscopic resection for upper gastrointestinal multiple submucosal tumors originating from the muscularis propria layer: A feasibility study*. Indian J Cancer, 2014. **51**: e52–e55. doi:10.4103/0019-509X.151989
14. Zhang X, Li X, Liu Y, Liu F. *The therapeutic effect of the digestive endoscopy tunneling technique on upper gastrointestinal muscularis propria tumors*. Am J Transl Res, 2021. **13**: 10663–10669.

15. Zhou DJ. *Submucosal tunneling and endoscopic resection of submucosal tumors at the esophagogastric junction*. World J Gastroenterol, 2015. **21**:578. doi:10.3748/wjg.v21.i2.578
16. Albrecht H, Raithel M, Braun A, Nagel A, Stegmaier A, Utpatel K, Schäfer C. *Endoscopic full-thickness resection (EFTR) in the lower gastrointestinal tract*. Tech Coloproctol, 2019. **23**:957–963. doi:10.1007/s10151-019-02043-5
17. Chen T, Zhang C, Yao LQ, Zhou PH, Zhong YS, Zhang YQ, Chen WF, Li QL, Cai MY, Chu Y, Xu MD. *Management of the complications of submucosal tunneling endoscopic resection for upper gastrointestinal submucosal tumors*. Endoscopy, 2016. **48**:149–155. doi:10.1055/s-0034-1393244
18. Ge N, Sun S, Wang S, Liu X, Wang G, Guo J. *Endoscopic ultrasound-assisted tunnel-type endoscopic submucosal dissection for the treatment of esophageal tumors arising in the muscularis propria (with video)*. Endosc Ultrasound, 2013. **2**:11–15. doi:10.7178/eus.04.004
19. Gong W, Xiong Y, Zhi F, Liu S, Wang A, Jiang B. *Preliminary experience of endoscopic submucosal tunnel dissection for upper gastrointestinal submucosal tumors*. Endoscopy, 2012. **44**:231–235. doi:10.1055/s-0031-1291720
20. He G, Wang J, Chen B, Xing X, Wang J, Chen J, Cui Y, Chen M. *Feasibility of endoscopic submucosal dissection for upper gastrointestinal submucosal tumors treatment and value of endoscopic ultrasonography in pre-operation assess and post-operation follow-up: a prospective study of 224 cases in a single medical center*. Surg Endosc, 2016. **30**:4206–4213. doi:10.1007/s00464-015-4729-1
21. Heresbach D, Kornhauser R, Seyrig JA, Coumaros D, Claviere C, Bury A, Cottureau J, Canard JM, Chaussade S, Baudet A, Casteur A, Duval O, Ponchon T; OMEGA group. *A national survey of endoscopic mucosal resection for superficial gastrointestinal neoplasia*. Endoscopy, 2010. **42**:806–813. doi:10.1055/s-0030-1255715
22. Inoue H, Ikeda H, Hosoya T, Onimaru M, Yoshida A, Eleftheriadis N, Maselli R, Kudo S. *Submucosal endoscopic tumor resection for subepithelial tumors in the esophagus and cardia*. Endoscopy, 2012. **44**:225–230. doi:10.1055/s-0031-1291659
23. Liu BR, Song JT, Kong LJ, Pei FH, Wang XH, Du YJ. *Tunneling endoscopic muscularis dissection for subepithelial tumors originating from the muscularis propria of the esophagus and gastric cardia*. Surg Endosc, 2013. **27**:4354–4359. doi:10.1007/s00464-013-3023-3
24. Lu J, Jiao T, Li Y, Liu Y, Wang Y, Wang Y, Zheng M, Lu X. *Heading toward the right direction - Solution package for endoscopic submucosal tunneling resection in the stomach*. PLoS One, 2015. **10**:1–9. doi:10.1371/journal.pone.0119870
25. Maehata T, Kato M, Ochiai Y, Mizutani M, Tsutsumi K, Kiguchi Y, Akimoto T, Sasaki M, Takatori Y, Nakayama A, Takabayashi K, Fujimoto A, Goto O, Yahagi N. *Feasibility of endoscopic submucosal dissection for colorectal neoplasia at anastomotic sites: a retrospective study*. Surg Endosc, 2020. **34**:5495–5500. doi:10.1007/s00464-019-07346-0

26. Meier B, Caca K, Schmidt A. *Hybrid endoscopic mucosal resection and full-thickness resection: a new approach for resection of large non-lifting colorectal adenomas (with video)*. Surg Endosc, 2017. **31**:4268–4274. doi:10.1007/s00464-017-5461-9
27. Nabi Z, Ramchandani M, Sayyed M, Darisetty S, Kotla R, Rao GV, Reddy DN. *Outcomes of submucosal tunneling endoscopic resection in upper gastrointestinal sub-epithelial tumors*. Indian J Gastroenterol, 2019. **38**:509–517. doi:10.1007/s12664-019-00988-x
28. Schmidt A, Bauerfeind P, Gubler C, Damm M, Bauder M, Caca K. *Endoscopic full-thickness resection in the colorectum with a novel over-the-scope device: First experience*. Endoscopy, 2015. **47**:719–725. doi:10.1055/s-0034-1391781
29. Song S, Feng M, Zhou H, Liu M, Sun M. *Submucosal tunneling endoscopic resection for large and irregular submucosal tumors originating from muscularis propria layer in upper gastrointestinal tract*. J Laparoendosc Adv Surg Tech, 2018. **28**:1364–1370. doi:10.1089/lap.2017.0607
30. Tan Y, Lv L, Duan T, Zhou J, Peng D, Tang Y, Liu D. *Comparison between submucosal tunneling endoscopic resection and video-assisted thoracoscopic surgery for large esophageal leiomyoma originating from the muscularis propria layer*. Surg Endosc, 2016. **30**:3121–3127. doi:10.1007/s00464-015-4567-1

