

ANSWERING REVIEWERS

September 25, 2015



Dear Editor,

Title: Second-look endoscopy and factors associated with delayed bleeding after endoscopic submucosal dissection

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Name of Journal: *World Journal of Gastrointestinal Endoscopy*

ESPS Manuscript NO: 18441

The manuscript has been improved according to the suggestions of reviewers:

1. Format has been updated

2. Revision has been made according to the suggestions of the reviewer

(1) First, bring the section of SLE AFTER ESD first, easy to read is better where the section of ROLE OF PROTON-PUMP INHIBITORS (PPIS) IN THE PREVENTION OF BLEEDING EVENTS behind.

Scoundary, you would not need to describe a method of searching the literature.

→ We have changed the order of contents as suggested

(2) Title Looked like that the author mainly described and discussed about post-ESD bleeding itself rather than second-look endoscopy. Please consider to revise the title.

→ As the reviewer pointed out, we have changed the title of this manuscript.

(3) There were some irreconcilable descriptions among the paragraphs and chapters. In addition, there were too many speculations through the manuscript.

Abstract

1, Gastric ESD is not noninvasive but also less invasive therapeutic endoscopy.

(page 3, lines 5-6)

Introduction

2, Circumferential mucosal incision is not always performed during gastric ESD, especially by needle-type device user. (page 5, lines 18-19)

3, Anemia is not cardiovascular adverse event. (page 6, line 1-2)

4, PPI and prophylactic coagulation of the mucosal defect has been widely spread and can be standard of care to reduce post-ESD bleeding. (page 6, line 4-5)

→ We have corrected as you suggested.

(4) Role of PPI to prevent post-ESD bleeding

The author should add the reference shown below and extend the discussion about the prevention of post-ESD bleeding. Uedo N, Takeuchi Y, Yamada T, Ishihara R, Ogiyama H, Yamamoto S, Kato M, Tatsumi K, Masuda E, Tamai C, Yamamoto S, Higashino K, Iishi H, Tatsuta M. Effect of a proton pump inhibitor or an H2-receptor antagonist on prevention of bleeding from ulcer after endoscopic submucosal dissection of early gastric cancer: a prospective randomized controlled trial. Am J Gastroenterol. 2007

→ We have added the reference (Uedo N et al Am J Gastroenterol 2007) and mentioned the results of meta-analysis and other recent studies about anti-secretory agents. (page 8, line 22 – page9, line 1)

(5) PROPHYLACTIC COAGULATION OF VISIBLE VESSELS

Please clarify which guidelines did the author mention about.

→ We have added the reference (Laine L, Jensen DM Am J Gastroenterol 2012, Hwang JH et al. Gastrointest Endosc 2012) as you suggested. (page 10, line 2)

(6) SLE

I don't think SLE still has been performed in many institutions, especially in Western countries which have outpatients-based medical system. Does the author have any data?

→ We agree with the reviewer about that. SLE is not performed in outpatients-based medical system. We have mentioned the ESD treatment setting (inpatient-based ESD treatment). (page 6, line 23-26)

(7) The author should introduce three prospective studies showing no clinical benefit of SLE to prevent post-ESD bleeding in detail and discuss the advantage and disadvantage of each study including study design and primary outcomes. The sample size was adequately calculated according to clinical hypothesis in some prospective studies.

→ We have described the primary outcomes and sample size calculation of three RCTs and discussed the disadvantage of each study. (page 7, line 4-25)

(8) DRUG-RELATED RISK FACTORS FOR DELAYED BLEEDING

Please consider to add the references below and extend the discussion.

Continuous aspirin use does not increase post-endoscopic dissection bleeding risk for gastric neoplasms in patients on antiplatelet therapy. Tounou S, Morita Y, Hosono T. *Endosc Int Open*. 2015 Feb;3(1):E31-8.

Continued use of low-dose aspirin does not increase the risk of bleeding during or after endoscopic submucosal dissection for early gastric cancer. Sanomura Y, Oka S, Tanaka S, Numata N, Higashiyama M, Kanao H, Yoshida S, Ueno Y, Chayama K. *Gastric Cancer*. 2014;17(3):489-96.

→ We have added the references as you suggested and discussed about the guidelines of ASGE, ESGE, Japan. (page 12, line 8-13 and line 20-23)

(9) the STRAP trial published by Dr. Ono probably included the patients undergoing other therapeutic endoscopy than gastric ESD.



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→ As the reviewer pointed out, the STRAP trial included the patients undergoing various therapeutic endoscopy. But the results mentioned in this manuscript is subanalysis of gastric ESD. We have added the comment about subanalysis of gastric ESD. (page 12, line 14)

Thank you again for publishing our manuscript in the *Gastrointestinal Endoscopy*.

Sincerely yours,

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