

PEER-REVIEW REPORT

Name of journal: *World Journal of Gastroenterology*

Manuscript NO: 81367

Title: Tranexamic acid may be a useful pharmacotherapy for endoscopically resistant small bowel angiodysplasia

Provenance and peer review: Unsolicited manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 06350183

Position: Peer Reviewer

Academic degree: MD

Professional title: Assistant Professor

Reviewer's Country/Territory: Romania

Author's Country/Territory: Japan

Manuscript submission date: 2022-11-05

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-11-06 15:38

Reviewer performed review: 2022-11-09 18:01

Review time: 3 Days and 2 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input type="checkbox"/> Yes <input checked="" type="checkbox"/> No

Peer-reviewer statements	Peer-Review: [<input checked="" type="checkbox"/>] Anonymous [<input type="checkbox"/>] Onymous
	Conflicts-of-Interest: [<input type="checkbox"/>] Yes [<input checked="" type="checkbox"/>] No

SPECIFIC COMMENTS TO AUTHORS

The manuscript is valuable as it highlights possible expanding of treatment approach from gastrointestinal vascular malformations in hereditary hemorrhagic telangiectasia (HHT) to small bowel angiodysplasia (SBAD) patients, based on similar clinical features, therapeutic methods and goals but also challenges. The manuscript addresses the key issue of hemorrhagic angiodysplasia in the small intestine that is difficult to treat endoscopically, bringing forward possible pharmacotherapy and presenting current knowledge and available trials results. The main hypothesis of the work is discussing evidence on the use of tranexamic acid in non-HHT patients suffering from bleeding gastrointestinal angiodysplasia, using HHT recent guidelines and relevant publications on the antifibrinolytic use of this drug. Maybe the author could consider if readership would value additional detailing on HHT guidelines (Faughnan ME, et al. Second International Guidelines for the Diagnosis and Management of Hereditary Hemorrhagic Telangiectasia. Ann Intern Med. 2020 Dec 15;173(12):989-1001. <https://doi.org/10.7326/M20-1443>) recommending that “clinicians consider treatment of mild HHT-related GI bleeding with oral antifibrinolytics” and “treatment of moderate to severe HHT-related GI bleeding with intravenous bevacizumab or other systemic antiangiogenic therapy”- so that similar approach would be evaluated in hemorrhagic angiodysplasia in the small bowel. The limitations of current knowledge are presented in a well-balanced manner, coming from scarcity of strong evidence and weak guidelines recommendations regarding specific use of tranexamic acid. Findings presented by the manuscript have the potential to significantly impact clinical practice in the direct benefit of patients suffering from difficult to manage GI hemorrhagic



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7041 Koll Center Parkway, Suite
160, Pleasanton, CA 94566, USA
Telephone: +1-925-399-1568
E-mail: bpgoffice@wjgnet.com
https://www.wjgnet.com

angiodysplasia.

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Peer-review model: Single blind

Reviewer's code: 03267595

Position: Peer Reviewer

Academic degree: PhD

Professional title: Professor

Reviewer's Country/Territory: China

Author's Country/Territory: Japan

Manuscript submission date: 2022-11-05

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-11-06 06:09

Reviewer performed review: 2022-11-11 07:12

Review time: 5 Days and 1 Hour

Scientific quality	<input checked="" type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input checked="" type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No



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https://www.wjgnet.com

Peer-reviewer statements	Peer-Review: [<input checked="" type="radio"/>] Anonymous [<input type="radio"/>] Onymous
	Conflicts-of-Interest: [<input type="radio"/>] Yes [<input checked="" type="radio"/>] No

SPECIFIC COMMENTS TO AUTHORS

Small bowel angiodysplasia (SBAD) is one of the common causes of gastrointestinal bleeding, and there are no recommended drugs to treat it now. The paper reports the pharmacological treatments of endoscopic difficult-to-treat SBAD, including the drug mechanism and side effects of them. The application of tranexamic acid in SBAD is also reported. Meanwhile, it is suggested to add more research reports in related fields to enhance the credibility and strength of the viewpoints. The paper was written concisely and clearly, providing drug therapy program support for patients with difficulty in endoscopic treatment of SBAD. The paper is of novel topic and is recommended to be accepted.

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Provenance and peer review: Unsolicited manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 05654229

Position: Peer Reviewer

Academic degree: MD

Professional title: Chief Physician, Professor

Reviewer's Country/Territory: China

Author's Country/Territory: Japan

Manuscript submission date: 2022-11-05

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-11-06 12:34

Reviewer performed review: 2022-11-12 03:23

Review time: 5 Days and 14 Hours

Scientific quality	<input checked="" type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input type="checkbox"/> Yes <input checked="" type="checkbox"/> No

Peer-reviewer statements	Peer-Review: [<input checked="" type="radio"/>] Anonymous [<input type="radio"/>] Onymous
	Conflicts-of-Interest: [<input type="radio"/>] Yes [<input checked="" type="radio"/>] No

SPECIFIC COMMENTS TO AUTHORS

First, tranexamic acid is recommended for patients with gastrointestinal angiodysplasias in hereditary hemorrhagic telangiectasia who are difficult to treat endoscopically. Investigation of the use of tranexamic acid for small intestinal angiodysplasia is desired. Second, there are several reports that hormone therapy, somatostatin analogs, thalidomide and VEGF-neutralizing antibodies are useful for SBAD for which endoscopic treatment is difficult. Third, although there are concerns about the risk of thrombosis and embolism, tranexamic acid is expected to reduce the amount of gastrointestinal bleeding in patients with SBAD in whom endoscopic treatment is difficult. Future reports are expected, as tranexamic acid could be a first-line drug for patients with SBAD.

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Peer-review model: Single blind

Reviewer's code: 03443948

Position: Peer Reviewer

Academic degree: PhD, MD

Professional title: Professor, Doctor

Reviewer's Country/Territory: China

Author's Country/Territory: Japan

Manuscript submission date: 2022-11-05

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-11-07 12:55

Reviewer performed review: 2022-11-13 15:16

Review time: 6 Days and 2 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
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	Conflicts-of-Interest: [<input type="radio"/>] Yes [<input checked="" type="radio"/>] No

SPECIFIC COMMENTS TO AUTHORS

Small bowel angiodysplasia is common disease and also common cause leading to small bowel bleeding. Indeed, the bleeding resulting from small bowel angiodysplasia is difficult to endoscopic treatment. This article reviews the drug treatment of small bowel angiodysplasia bleeding, especially reviewing the efficacy and safety of tranexamic acid, a traditional antifibrinolytic agent, which provides a valuable reference for treatment of small bowel angiodysplasia bleeding. I think it can be published in this journal. My question: 1.The title is long, and did not reflect the main subject of the manuscript, suggesting revision. 2.English is not good, needing improvement.