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PEER-REVIEW REPORT

Name of journal: *World Journal of Gastrointestinal Endoscopy*

Manuscript NO: 85689

Title: Polyposis found on index colonoscopy in a 56-year-old female - BMP YA variant in juvenile polyposis syndrome: A case report

Provenance and peer review: Unsolicited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 03210228
Position: Peer Reviewer
Academic degree: MD

Professional title: Professor

Reviewer's Country/Territory: China

Author's Country/Territory: Australia

Manuscript submission date: 2023-05-10

Reviewer chosen by: Geng-Long Liu

Reviewer accepted review: 2023-05-29 02:35

Reviewer performed review: 2023-06-05 06:08

Review time: 7 Days and 3 Hours

Scientific quality	[ ] Grade A: Excellent [Y] Grade B: Very good [ ] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No novelty
Creativity or innovation of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No creativity or innovation

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Scientific significance of the conclusion in this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No scientific significance
Language quality	[ ] Grade A: Priority publishing [Y] Grade B: Minor language polishing [ ] Grade C: A great deal of language polishing [ ] Grade D: Rejection
Conclusion	[ ] Accept (High priority) [Y] Accept (General priority) [ ] Minor revision [ ] Major revision [ ] Rejection
Re-review	[Y] Yes [ ] No
Peer-reviewer statements	Peer-Review: [ ] Anonymous [Y] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

This manuscript provides a comprehensive description of the diagnosis and management of an asymptomatic JPS patient. Through this case, the authors reviewed the management suggestions for such patients in the guidelines of the US and Europe, and provided appropriate measures according to the patient's actual condition. The authors emphasizes several important issues. 1. The differential diagnosis of polyposis subtypes is very difficult, and it is not mandatory for clinical doctors. But at the same time, for gastroenterologists, the key is to identify it, that is, to have a clear understanding of the intestinal and extraintestinal manifestations of polyposis, so that the subsequent treatments can be provided. 3. Benefiting from the development of NGS, the differential diagnosis of polyposis can be accomplished with the help of gene test and genetic counseling. Through this fortunate case of early diagnosis and treatment, we will understand more about JPS. The authors can also further provide readers with information on the phenotypic complexity of BMPR1A associated JPS based on this case. Please refer to this article (PMID 36632626), which showed the phenotype of BMPR1A associated diseases can range from colorectal cancer without polyps to polyposis with

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more than 100 polyps like in this case. The phenotypic complexity was discovered with the help of the widespread use of NGS. More important, the readers should also recognize that the discovery of genetic variations maybe earlier than the emergence of classic phenotypes in these patients, and these new insights will help readers provide more accurate guidance for patients. The 'ultra rare' in the title should be replaced by 'reported'. For rare diseases, it is strange to emphasize the rarity of pathogenic variants, and it is customary to express them in the binary form of 'novel' and 'reported'. There can be high-frequency mutations in expression to correspond to mutation hotspots, but the opposite situation has not been reported in the literature.

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Peer-review model: Single blind

Reviewer's code: 02451447
Position: Editorial Board
Academic degree: MD, PhD

Professional title: Professor

Reviewer's Country/Territory: United States

Author's Country/Territory: Australia

Manuscript submission date: 2023-05-10

Reviewer chosen by: Geng-Long Liu

Reviewer accepted review: 2023-06-27 02:17

Reviewer performed review: 2023-06-27 02:36

Review time: 1 Hour

Scientific quality	[ ] Grade A: Excellent [ ] Grade B: Very good [Y] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No novelty
Creativity or innovation of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No creativity or innovation

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Scientific significance of the conclusion in this manuscript	[ ] Grade A: Excellent [ ] Grade B: Good [Y] Grade C: Fair [ ] Grade D: No scientific significance
Language quality	[ ] Grade A: Priority publishing [Y] Grade B: Minor language polishing [ ] Grade C: A great deal of language polishing [ ] Grade D: Rejection
Conclusion	[ ] Accept (High priority) [ ] Accept (General priority) [Y] Minor revision [ ] Major revision [ ] Rejection
Re-review	[ ] Yes [Y] No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

The authors reported a juvenile polyposis syndrome case with BMPR1A mutation. Comment: 1. I would not recommend to the words such as "extensive polyposis" and "ultra-rare" in the title and the entire manuscript. Please remove "extensive" and "ultra". 2. In the abstract, please do not give very basic information in the background, such as the incidence rate of JPS. This information should be present in the introduction or discussion. Please try to keep the abstract more concise. 3. I would suggest to list "Cronkhite-Canada syndrome " as a differential diagnosis, since the morphology and clinical presentation do not fit Cronkhite-Canada syndrome.

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Peer-review model: Single blind

Reviewer's code: 03544596
Position: Editorial Board
Academic degree: MD
Professional title: Academic Editor, Associate Professor
Reviewer's Country/Territory: Turkey
Author's Country/Territory: Australia

Manuscript submission date: 2023-05-10

Reviewer chosen by: Geng-Long Liu

Reviewer accepted review: 2023-06-27 07:27

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Review time: 1 Hour

Scientific quality	[ ] Grade A: Excellent [Y] Grade B: Very good [ ] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No novelty
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Conclusion	[ ] Accept (High priority) [Y] Accept (General priority) [ ] Minor revision [ ] Major revision [ ] Rejection
Re-review	[ ] Yes [Y] No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

Dear Editor, I should first thank for inviting me as potential reviewer to read and comment on paper entitled ''Extensive polyposis found on index colonoscopy in a 56-year-old female: A case report on an ultra-rare BMPR1A variant in juvenile polyposis syndrome''. In the current study, the authors aimed to report a case of extensive polyposis found on index screening endoscopy in an asymptomatic female with no prior related family or medical history. The main title accurately reflects the major topic and content of the study. The abstract summarizes and reflects the work described in the manuscript. Also, the abstract presents the significant points related to the background, objectives, materials and methods, results and conclusions. The section of the discussion is well organized. The conclusions are drawn appropriately supported by the literature. The manuscript adequately describes the background, present status and significance of the study. The manuscript interprets the findings adequately and appropriately, highlighting the key points clearly. I think that it will contribute to the literature.

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Provenance and peer review: Unsolicited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 02954019
Position: Editorial Board
Academic degree: MD, PhD
Professional title: Associate Professor
Reviewer's Country/Territory: Japan
Author's Country/Territory: Australia

Manuscript submission date: 2023-05-10

Reviewer chosen by: Geng-Long Liu

Reviewer accepted review: 2023-06-27 07:05

Reviewer performed review: 2023-06-27 07:52

Review time: 1 Hour

Scientific quality	[ ] Grade A: Excellent [ ] Grade B: Very good [ ] Grade C: Good [Y] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[ ] Grade A: Excellent [ ] Grade B: Good [ ] Grade C: Fair [Y] Grade D: No novelty
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Re-review	[ ] Yes [Y] No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

This is a reported case of juvenile polyposis (JPS) without the commonly known pathological variant of the BMPR1A gene. On the other hand, about 70% of JPS without pathological variants of the BMPR1A gene have been reported. This can be interpreted as many cases do not have a pathological variant. It is inconsistent to assume that the presence of the BMPR1A c.1409T>C (p.Met470Thr) mutation, which has been associated with JPS in only two cases so far, including the present report, is associated with JPS. Without evidence that the c.1409T>C (p.Met470Thr) mutation is present in a certain percentage of JPS cases, we consider this reported case to be of little significance.