

PEER-REVIEW REPORT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 01587889

Position: Editorial Board

Academic degree: MD, MSc, PhD

Professional title: Associate Professor, Lecturer, Surgeon

Reviewer's Country/Territory: United States

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: Andrea Lisotti

Reviewer accepted review: 2021-06-12 13:33

Reviewer performed review: 2021-06-12 13:53

Review time: 1 Hour

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 Grade E: Do not publish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
Conclusion	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input checked="" type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input 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



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SPECIFIC COMMENTS TO AUTHORS

Please, advise the corresponding author to adjust the title of the paper reflecting its contents. I suggest the title should read *Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis*. The findings reported in the paper that the dietary modifications and probiotics seem to confer the greatest benefit for pouch-related symptoms than antibiotics and anti-inflammatory modalities is very reasonable.

PEER-REVIEW REPORT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 00536689

Position: Editorial Board

Academic degree: MD

Professional title: Assistant Professor, Surgeon

Reviewer's Country/Territory: Italy

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: Andrea Lisotti

Reviewer accepted review: 2021-06-14 10:54

Reviewer performed review: 2021-06-14 11:03

Review time: 1 Hour

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input checked="" 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 type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input checked="" type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No
Peer-reviewer statements	Peer-Review: <input type="checkbox"/> Anonymous <input checked="" type="checkbox"/> Onymous Conflicts-of-Interest: <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No



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SPECIFIC COMMENTS TO AUTHORS

I have read the manuscript carefully. Current topic. Usually, ileal pouches packaged for familial polyposis have a lower incidence of pouchitis than patients operated on for ulcerative colitis. Comprehensive introduction. There is no information on the type of ileal pouch. No information is reported on whether patients had a protective ileostomy. No information on ileostomy closure. I think the authors should clarify these aspects. Further information on the surgical act is absent.

PEER-REVIEW REPORT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 03733992

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: United States

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: AI Technique

Reviewer accepted review: 2021-06-13 22:49

Reviewer performed review: 2021-06-14 16:26

Review time: 17 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input checked="" 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 type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input checked="" type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input 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



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SPECIFIC COMMENTS TO AUTHORS

Thank you for allowing me to review this article. It is indeed in any unexplored area in which research is certainly needed especially as it greatly impacts quality of life for patients with FAP who have had a TPC-IPAA. The study is majorly flawed, however, and there are multiple issues that need to be addressed before it can be considered for publication. I am happy to re-review once this has been done. - I do not fault the authors for a study with 33 individuals since FAP is rare and such studies understandably will have a small overall n. That being said, and though it is understandably retrospective, the numbers for each intervention are too small to make conclusions. The statistics of this will invariably influence readers to think that certain interventions are successful but may instead be due to chance alone. I think all interventions can only be presented as "grouped together." Though a table can be provided with number treated by each intervention, conclusions of the effect of each should not be teased out. -The pathophysiology of pouchitis in IBD is completely different than that in FAP. IBD is driven by an inflammatory process, while as the authors astutely mention, FAP pouch issues likely by an IBS-related process. The PDAI, developed for IBD, thus cannot be used in this study. Instead, the authors should evaluate symptoms individually. This is further supported by the fact that only n=4 had overt pouchitis and there was minimal non-significant response to anti-inflammatory therapies. -On a related note, there are no tables (not sure if I didnt receive any but I dont see a supplemental document). I suspect that the greatest impact from the PDAI comes from number of bowel movements. Again, symptoms should be described individually and PDAI is inappropriate to use here. -The lack of a placebo control group as well as inability to describe why each intervention was used makes this a descriptive study. -This study comes from a tertiary referral center. How many



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professionals made medical decisions to treat these patients and pick individual interventions? How were medications chosen? More details should be provided - was there an algorithm, was it provider specific, etc? -The difference between statistical significance and clinical significance should be emphasized and elaborated upon. A decrease in DBM from 10.3 to 9.3 is clinically insignificant even though it reaches statistical significance. If readers conclude that interventions will significantly reduce bowel movements, the point will be lost. A patient with 9 BMs is no different than one with 10. -For the patients with overt pouchitis, did any have IBD?

PEER-REVIEW REPORT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 03805385

Position: Peer Reviewer

Academic degree: FASCRS, MD, PhD

Professional title: Assistant Professor, Attending Doctor, Doctor, Medical Assistant, Postdoctoral Fellow, Research Associate, Senior Research Fellow, Surgeon

Reviewer's Country/Territory: Brazil

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: Andrea Lisotti

Reviewer accepted review: 2021-06-13 15:42

Reviewer performed review: 2021-06-15 08:32

Review time: 1 Day and 16 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input checked="" 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 type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input checked="" type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No
Peer-reviewer	Peer-Review: <input checked="" type="checkbox"/> Anonymous <input type="checkbox"/> Onymous

statements

Conflicts-of-Interest: [] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

Thank you very much for the opportunity to review this paper. Major comments: The title, abstract and the key words reflect the study described by the authors. I have some trouble to understand the Methods. You need to describe data analysis in detail, and how do you make your analysis. On the other hand, the authors referred to the tables 1 and 2, but I never found the tables in your manuscript. Do you compared the different types of treatment? Do you only analyzed the outcomes of every treatment compared by themselves? The results are very difficult to follow. A diagram of the different treatment need to be created in the methods. You have a good manuscript, with good information but the information need to improved. Finally it is important that you stated: this is an retrospective work.

PEER-REVIEW REPORT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 06117003

Position: Peer Reviewer

Academic degree: MS

Professional title: Professor

Reviewer's Country/Territory: India

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: AI Technique

Reviewer accepted review: 2021-06-13 07:33

Reviewer performed review: 2021-06-21 15:50

Review time: 8 Days and 8 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input checked="" 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



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SPECIFIC COMMENTS TO AUTHORS

The manuscript addresses an important aspect in the management of IPS in a rare subset of patients with APS who have undergone IPAA. The authors have nicely elaborated the issue and have suggested some definitive management strategies. The authors have extrapolated their work retrospectively from their own prospective observational study which was a “by chance” finding. It was perhaps an accidental result rather than a pre-planned study with definite parameters. Though the results and statistical analysis favours some definitive management strategies based on treatment according to lines of IBS, a well-defined cohort based prospective study could help in generating definitive treatment protocol. Though the study opens up a new avenue in the management of IPS but acceptance of treatment based on the conclusions of the Author’s needs further research before being implemented into clinical practise.

PEER-REVIEW REPORT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 02541859

Position: Editor-in-Chief

Academic degree: FACG, FACP, FASGE, FRCP, MD, MRCP

Professional title: Associate Professor, Doctor

Reviewer's Country/Territory: United States

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: AI Technique

Reviewer accepted review: 2021-06-14 17:58

Reviewer performed review: 2021-06-22 11:53

Review time: 7 Days and 17 Hours

Scientific quality	<input 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 checked="" type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input checked="" type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input checked="" 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 study has mentioned about dietary modifications but what kind of dietary modification was done is not elaborated. It is difficult to establish that just symptomatic treatment or dietary modifications improved patients' symptoms. The PDAI should have been tabulated to compare different treatment groups.

PEER-REVIEW REPORT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 03478442

Position: Peer Reviewer

Academic degree: FEBG, MD, PhD

Professional title: Professor

Reviewer's Country/Territory: Greece

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: Andrea Lisotti

Reviewer accepted review: 2021-06-18 07:49

Reviewer performed review: 2021-06-23 21:37

Review time: 5 Days and 13 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
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 checked="" type="checkbox"/> Yes <input 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



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SPECIFIC COMMENTS TO AUTHORS

it is a very interesting and well written manuscript, presenting data about management of pouch related symptoms in patients with adenomatous polyposis. However, this study has several limitations, such as the small number of patients and only four patients with identified pouchitis (>7 PDAI). However, data about this subgroup of patients with ileal-pouch anal anastomosis are very limited. Furthermore the authors analyze their findings and compare them with current literature and this is an attractive feature of the manuscript.

PEER-REVIEW REPORT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 04718191

Position: Peer Reviewer

Academic degree: MD, PhD

Professional title: Professor, Surgeon

Reviewer's Country/Territory: China

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: Andrea Lisotti

Reviewer accepted review: 2021-06-13 12:19

Reviewer performed review: 2021-06-26 14:28

Review time: 13 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
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 checked="" type="checkbox"/> Yes <input 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 authors assessed clinical, endoscopic and histologic response to various treatments in the pouch related disorders of APS patients. Thirty-three APS patients were identified. Intervention was associated with symptomatic relief, mainly decreasing abdominal pain and daily bowel movements. Dietary modifications decreased abdominal pain, daily bowel movements, overall PDAI and clinical PDAI. Probiotics decreased daily bowel movements, overall and clinical PDAI histologic scores. The authors suggested pouch-related symptoms a functional rather than inflammatory disorder. I have some comments: 1. How to define pouch related symptoms? Bloody stools usually indicated an inflammatory problem. It could be better to exclude the patients with pouchitis in the cohort, while only analyse the patients with IPS. 2. I cannot see the tables and figures in the Manuscript file. 3. As you stated that average 3 different therapies were used per patient, how can you separate the effects of individual therapies from another? 4. In the conclusion, the authors suggested pouch-related symptoms in APS a functional rather than inflammatory disorder, again you should separate pouchitis from IPS.

RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 03805385

Position: Peer Reviewer

Academic degree: FASCRS, MD, PhD

Professional title: Assistant Professor, Attending Doctor, Doctor, Medical Assistant, Postdoctoral Fellow, Research Associate, Senior Research Fellow, Surgeon

Reviewer's Country/Territory: Brazil

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: Jin-Lei Wang

Reviewer accepted review: 2021-07-29 22:15

Reviewer performed review: 2021-07-30 19:01

Review time: 20 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 checked="" type="checkbox"/> Grade A: Priority publishing <input 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
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



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SPECIFIC COMMENTS TO AUTHORS

The authors response every question, I believe the journal can publish this paper, if the others reviewers agreed. Thanks again for the opportunity. As I said before this paper has an excellent quality and the conclusion was appropriate. And the limitations were improved The title reflect the subject of the manuscript The abstract summarize and reflect the work in the manuscript The key words reflect the focus of the manuscript

RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 68214

Title: Management of pouch related symptoms in patients who underwent ileal pouch anal anastomosis surgery for adenomatous polyposis

Reviewer's code: 00536689

Position: Editorial Board

Academic degree: MD

Professional title: Assistant Professor, Surgeon

Reviewer's Country/Territory: Italy

Author's Country/Territory: Israel

Manuscript submission date: 2021-06-12

Reviewer chosen by: Jin-Lei Wang

Reviewer accepted review: 2021-07-29 07:33

Reviewer performed review: 2021-08-01 07:17

Review time: 2 Days and 23 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 checked="" type="checkbox"/> Grade A: Priority publishing <input 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
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

I recommend to accept the paper.