

**ESPS Peer-review Report**
**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 02548813

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2013-10-02 02:20

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A (Excellent)	<input type="checkbox"/> Grade A: Priority Publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B (Very good)	<input checked="" type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<input type="checkbox"/> High priority for publication
<input checked="" type="checkbox"/> Grade C (Good)	<input type="checkbox"/> Grade C: a great deal of language polishing	<input type="checkbox"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D (Fair)	<input type="checkbox"/> Grade D: rejected	BPG Search:	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E (Poor)		<input type="checkbox"/> Existed	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

**COMMENTS TO AUTHORS**

The manuscript deals an interesting topic regard microbiota dysbiosis and pouchitis. The authors chose the main relevant articles to include in the manuscript, revealing some that show the same result and others, conflicting findings regarding the topic. It is a well written article, and I listed below some suggestions and considerations.

- Regarding the phrase “the presence of Clostridiaceae spp adherent to pouch mucosa and the high expression of TLR-2 and TLR-4 mRNA were predictive of chronic/relapsing pouchitis.” I suggest, for instance, “the presence of Clostridiaceae spp adherent to pouch mucosa and the high expression of TLR-2 and TLR-4 mRNA were associated to chronic/relapsing pouchitis.”, and not they predict this complication (last paragraph before the conclusions). We usually say that a molecule or a factor predicts chronic or recurrence disease when it has been tested in an independent cohort, and in a longitudinal study.
- Restorative proctocolectomy (RPC) and restorative proctocolectomy with ileal pouch-anal anastomosis (IPAA) are the same procedures, so I suggest chose one of the nomenclature (IPAA or RPC).
- Please, review the following references in accordance to the journal rules:
  - Garrett C. Zella, MD,\* Elizabeth J. Hait, MD, MPH,? Tiffany Glavan, BA,? Dirk Gevers, PhD,§,? Doyle V. Ward, PhD,k Christopher L. Kitts, PhD,\*\* and Joshua R. Korzenik, MD?? Distinct Microbiome in Pouchitis Compared to Healthy Pouches in Ulcerative Colitis and Familial Adenomatous Polyposis. *Inflamm Bowel Dis* ! 2011, Volume 17, Number 5, 1092, 1100.
  - Gerald W. Tannock, PhD,\* Blair Lawley, PhD,\* Karen Munro,\* Christophe Lay, PhD,\* Corinda Taylor,\* Christopher Daynes, MB, CHB,? Lori Baladjay, BSc,? Robin Mcleod, MD,? and Mark Thompson-Fawcett, MD? Comprehensive Analysis



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of the Bacterial Content of Stool from Patients with Chronic Pouchitis, Normal Pouches, or Familial Adenomatous Polyposis Pouches. Inflamm Bowel Dis ! Volume 18, Number 5, 2012; 925, 934 4.  
Quality of written English: proper.

**ESPS Peer-review Report**
**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 00723963

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2013-10-15 00:20

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A (Excellent)	<input type="checkbox"/> Grade A: Priority Publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B (Very good)	<input type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<input type="checkbox"/> High priority for publication
<input checked="" type="checkbox"/> Grade C (Good)	<input checked="" type="checkbox"/> Grade C: a great deal of language polishing	<input type="checkbox"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D (Fair)	<input type="checkbox"/> Grade D: rejected	<input type="checkbox"/> Existed	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E (Poor)		<input type="checkbox"/> No records	<input checked="" type="checkbox"/> Major revision

**COMMENTS TO AUTHORS**

This manuscript deals with an interesting topic and one which is a real clinical problem. The topic of pouchitis and its aetiology in terms of microbiota is an interesting one for both gastroenterologists and colorectal surgeons. I have the following comments: 1. My main criticism with regard to the manuscript is that it is a narrative review and not a systematic review and i wonder why the authors choose this format. A systematic review with a study flow diagram and an outline of how many articles were found from the initial search, how many and why were certain articles excluded etc is all standard information that is given in a systematic review and gives a better overall view of the literature to a reader. 2.The last 9 lines of the introduction is really the "search strategy" which should be part of the methodology section of a systematic review. Minor comments: Multiple grammatical errors including a. Page 6 1/2 way down "In lane with previous reports..." should be "In line with previous reports" b. Page 7 second paragraph: "associated to chronic/ relapsing pouchitis..." should be "associated with chronic/ relapsing pouchitis..." c. Page 7 1/2 way down: "These genes activation results...." should read "This genetic activation results...." d. Page 8 second paragraph: "There are evidences suggesting ...." should read "There is evidence suggesting...."

**ESPS Peer-review Report**

**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 00036648

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2014-02-09 06:21

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input checked="" type="checkbox"/> Grade A (Excellent)	<input checked="" type="checkbox"/> Grade A: Priority Publishing	Google Search:	<input checked="" type="checkbox"/> Accept
<input type="checkbox"/> Grade B (Very good)	<input type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C (Good)	<input type="checkbox"/> Grade C: a great deal of	<input type="checkbox"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D (Fair)	language polishing	BPG Search:	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E (Poor)	<input type="checkbox"/> Grade D: rejected	<input type="checkbox"/> Existed	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

**COMMENTS TO AUTHORS**

This is an interesting topic for review and the manuscript is well written.

**ESPS Peer-review Report**

**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 00503404

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2014-02-09 17:51

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A (Excellent)	<input type="checkbox"/> Grade A: Priority Publishing	Google Search:	<input type="checkbox"/> Accept
<input checked="" type="checkbox"/> Grade B (Very good)	<input checked="" type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C (Good)	<input type="checkbox"/> Grade C: a great deal of	<input type="checkbox"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D (Fair)	language polishing	BPG Search:	<input checked="" type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E (Poor)	<input type="checkbox"/> Grade D: rejected	<input type="checkbox"/> Existed	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

**COMMENTS TO AUTHORS**

A comprehensive review on the importance of microbial factors in the development of pouchitis  
Comments; 1. Please rename patients and methods to search strategy 2. Include an additional Table on how authors have identified the studies included

**ESPS Peer-review Report**
**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 00038879

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2014-02-11 06:38

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
[ Y] Grade A (Excellent)	[ ] Grade A: Priority Publishing	Google Search:	[ ] Accept
[ ] Grade B (Very good)	[ Y] Grade B: minor language polishing	[ ] Existed	[ ] High priority for publication
[ ] Grade C (Good)	[ ] Grade C: a great deal of language polishing	[ ] No records	[ ] Rejection
[ ] Grade D (Fair)	[ ] Grade D: rejected	[ ] Existed	[ Y] Minor revision
[ ] Grade E (Poor)		[ ] No records	[ ] Major revision

**COMMENTS TO AUTHORS**

1. there are numerous errors of language in tenses, plurals, wording, etc. This needs to be carefully edited by a native English speaker. 2. There are several instances where acronyms are either not defined or are defined later than first use, such as with RPC, TRFLP, OTU, CD, NP, CP-off, and CP-on. In addition, the authors use RPC, IPAA, pouch, ileo-anal pouch, ileal pouch, etc. interchangeably. The authors should pick one and be consistent. 3. There are confusing or poorly worded sentences. The definition of pouchitis by Mahadevan and Sandborn can be clarified. In the methods section, second sentence, the word "affection" is likely wrong. It makes the intent of the authors unclear. Similarly, in the sentence beginning "A manual cross reference..." the meaning of "originally resulting compatible" is unclear. 4. The table, as presented to the reviewer, was poorly constructed and difficult to read. Perhaps a landscape format, instead of letter format, as well as a smaller font, would make this more clear. 5. The title speaks specifically to the mucosa-associated microbiota. However, the majority of the review is on luminal microbiota. If it is the intent of the authors (and I believe it was) to emphasize this point, then the section on luminal microbiota can be shortened and more emphasis on mucosa-associated microbiota be placed. And along this line, the conclusion does not specifically address this issue as stated in the title. If it was not the authors' intention to differentiate the mucosa associated microbiota from luminal microbiota, then the title should be adjusted accordingly. Overall, I believe the authors should make a major effort to shorten this paper - it is way too long

**ESPS Peer-review Report**
**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 00180872

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2014-02-11 23:27

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A (Excellent)	<input type="checkbox"/> Grade A: Priority Publishing	Google Search:	<input checked="" type="checkbox"/> Accept
<input checked="" type="checkbox"/> Grade B (Very good)	<input checked="" type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C (Good)	<input type="checkbox"/> Grade C: a great deal of	<input type="checkbox"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D (Fair)	language polishing	BPG Search:	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E (Poor)	<input type="checkbox"/> Grade D: rejected	<input type="checkbox"/> Existed	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

**COMMENTS TO AUTHORS**

Thank you for the opportunity of reviewing this interesting manuscript. It is indeed a comprehensive review on pouch bacteriology. It is nicely written yet it is not easy to read; the paragraphs are somewhat long and sometimes repetitive. Try breaking the text to shorter paragraphs. Figure 1 is not available for review.

**ESPS Peer-review Report**
**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 00503405

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2014-02-15 14:28

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A (Excellent)	<input checked="" type="checkbox"/> Grade A: Priority Publishing	Google Search:	<input checked="" type="checkbox"/> Accept
<input checked="" type="checkbox"/> Grade B (Very good)	<input type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C (Good)	<input type="checkbox"/> Grade C: a great deal of	<input type="checkbox"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D (Fair)	language polishing	BPG Search:	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E (Poor)	<input type="checkbox"/> Grade D: rejected	<input type="checkbox"/> Existed	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

**COMMENTS TO AUTHORS**

The authors made a well written, interesting review about pouch bacteriota and its pathogenic role. The used references are up-to-date. Though I was not able to open Figure 1, the content of the manuscript is such good that I suggest to accept it for publication in WJG.



**ESPS Peer-review Report**
**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 00503587

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2014-02-16 06:35

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A (Excellent)	<input type="checkbox"/> Grade A: Priority Publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B (Very good)	<input checked="" type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<input type="checkbox"/> High priority for publication
<input checked="" type="checkbox"/> Grade C (Good)	<input type="checkbox"/> Grade C: a great deal of language polishing	<input type="checkbox"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D (Fair)	<input type="checkbox"/> Grade D: rejected	<input type="checkbox"/> Existed	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E (Poor)		<input type="checkbox"/> No records	<input checked="" type="checkbox"/> Major revision

**COMMENTS TO AUTHORS**

This review examines the impact of the flora and host responses in pouchitis. Specific Comments: 1. There are a number of errors of English grammar and/or construction that require correction. 2. In the Abstract, what is meant by RPC? Abbreviation should be explained here/in intro. 3. Another reason for "pouchitis" is that the disease is actually CD, not UC. 4. In the Methods, what is meant by pouch affection? 5. With regards to the flora and pouchitis, the authors could expand a little more on the impact of pouchitis therapies upon the flora. 6. What is meant by "chronic pouchitis off"? (page 8) 7. The Table needs to be reformatted so that words are complete.

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**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 00227376

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2014-02-24 06:48

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A (Excellent)	<input type="checkbox"/> Grade A: Priority Publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B (Very good)	<input checked="" type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<input type="checkbox"/> High priority for publication
<input checked="" type="checkbox"/> Grade C (Good)	<input type="checkbox"/> Grade C: a great deal of language polishing	<input type="checkbox"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D (Fair)	<input type="checkbox"/> Grade D: rejected	<input type="checkbox"/> Existed	<input checked="" type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E (Poor)		<input type="checkbox"/> No records	<input type="checkbox"/> Major revision

**COMMENTS TO AUTHORS**

This paper is a review of studies showing that dysbiosis of the ileal pouch microbiota has a role in the pathogenesis of pouchitis by impairing the regulation of mucosal immune system. In the paper, intestinal microbiota is classified into 2 groups: mucosal-adherent microbiota and luminal microbiota. Although the authors mentioned on these in the manuscript, there is a need for an in-depth discussion of these 2 concepts of microbiota. The authors should refer to the methods of culturing and showing the presence of these 2 groups of microbiota. Methodological differences between the published literature and the ideal method should be discussed in the manuscript. Also, it should be noted if mucosal-adherent microbiota can be affected by the treatment with antibiotics. Is microbiota of patients with refractory pouchitis different from the patients with responsive? Are there any evidence showing this possible difference is related to mucosal-adherent microbiota? With the clarification of these questions, the results of the study will become more apprehensible and efficient.

**ESPS Peer-review Report**
**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 6035

**Title:** Relationship between mucosa-associated microbiota and pouchitis following restorative proctocolectomy for ulcerative colitis

**Reviewer code:** 00503628

**Science editor:** Ma, Ya-Juan

**Date sent for review:** 2013-10-01 21:14

**Date reviewed:** 2014-02-24 23:50

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A (Excellent)	<input type="checkbox"/> Grade A: Priority Publishing	Google Search:	<input checked="" type="checkbox"/> Accept
<input checked="" type="checkbox"/> Grade B (Very good)	<input checked="" type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C (Good)	<input type="checkbox"/> Grade C: a great deal of	<input type="checkbox"/> No records	
<input type="checkbox"/> Grade D (Fair)	language polishing	BPG Search:	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade E (Poor)	<input type="checkbox"/> Grade D: rejected	<input type="checkbox"/> Existed	<input type="checkbox"/> Minor revision
		<input type="checkbox"/> No records	<input type="checkbox"/> Major revision

**COMMENTS TO AUTHORS**

This is a succinct well written and informative review on the analysis of the microbiota in the ileal anal pouch after ulcerative colitis and FAP. The subject matter is of high interest and is timely. It provides state of the art information on the subject. However, the table that is included is difficult to assess due to the format in which it was presented. Some spelling revisions may be needed. For instance, the last name of an author referred to in the text and in the table is spelled differently.