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**Name of Journal:** *World Journal of Transplantation*

**ESPS Manuscript NO:** 22906

**Manuscript Type:** REVIEW

### **Response to reviewers**

#### *First reviewer*

This paper is a fairly comprehensive review on the subject with extensive literature collected and reviewed. The overall write up of the subject is excellent, w/ only minor typos noted. Good review on historical background, as well. The current technology in general not able to develop functional unit of gut, in terms of functional secretion and interaction on peritoneal surface, venous and lymphatic drainage, muscular layer and neural innervations, as well as maintaining anti-bacterial barrier function. In doing so, the gut has not only a three-dimensional structure, but also an almost “four-dimensional” feature of individual subunits and layer, functionality and interaction with time/mobility Some reference should be made to peritoneal fluid as a rich source of stem cells Needs some upfront explanation in test: -page 9., 3rd para: explain word “nitrergic” -page 9., line before last: explain “organoids” upfront Typos: -page 4, 3rd line from last: the part “...procedure since however, SEP...” appears to have editing problem -page 5, line 4-5: Sentence starting as: “However, Bianchi procedure was...” - is confusing, unclear, what the writer intends to say. Please, rephrase to clarify statement. Additional, next sentence, needs a comma after “In addition” -Page 5, line before last - word “Supporting” has a come after it, appears not needed -page 8, mid-part of first para: “SIS” stand for “small intestinal sub mucosa” -- please correct -page 11, paragraph before last/ 6th line before last: correct typo in word “mascularis”

1. In our search in the literature we were not able to find any references regarding the use of peritoneal stem cells for intestinal regeneration.
2. Nitrergic neuronal population explained as Nerve cells where transmission is mediated by nitric oxide.

3. Organoids explained as “Then Vacanti and Choi, developed a villus structure with a core of mesenchymal stromal cells overlaid with epithelium called “Organoid Unit”.
4. Page 4, 3rd line from last changed to: However the patients undergone Bianchi procedure would wean off TPN more than those with STEP, they eventually would require intestinal transplants more than those with STEP.
5. A comma added after “In addition,”
6. A come after “Supporting” deleted.
7. Page 8, mid-part of first para: “SIS” stand for “small intestinal sub mucosa” corrected.
8. Page 11, paragraph before last/ 6th line before last: corrected typo in word “masclaris” to muscularis.

*Second reviewer*

The authors describe all the approaches to increase intestinal length, including intestinal regeneration and bioengineering. The manuscript is interesting, complete and well written. All the limitations are described and future directions are highlighted in this complex field.

No comments to address.