

Serial number	Reviewer comments	Addressed or rebuttal
1.	(Reviewed by 00504316): Basically, this is a good paper which well demonstrated the crosstalk between intra-islet endothelial cells and β -cells. Schematic photography is appreciated if possible to make the crosstalk relationship more clear	1. addressed
2.	(Reviewed by 00070310): This paper reviews small implication for islet cell transplantation. This paper is interest, and well written and appropriate. It will be acceptable for publication except one. Please show how to select papers in this review.	1. addressed
3.	(Reviewed by 00503542): This review article focuses on crosstalk between intra-islet endothelial cells and beta-cells. Recent advances are concisely summarized and informative for readers. However, a few small problems may remain to be solved as follows. 1. A section of "The endocrine effect of islet ECs on bet-cells" (page 5) may need reference(s). 2. In "when co-transplanted (23) or coated (24) with ECs in diabetes induced mice," (page 3), reference #23 seems to use diabetic rats. 3. In "(KDR in humans and Flk-1 in mouse)" (page 4), "mouse" may be written in plural form. 4. The last reference (#173) does not seem to have relevance to islets. If any, it should be mentioned. 5. Careful proof reading may improve some details. For example, "Beta-cell" is found on page 3. In the title of reference 24 (page11), the first letter of a word is capitalized.	1. addressed 2.addressed 3.addressed 4.addressed 5. addressed
4.	(Reviewed by 02993037): This is an interesting review which explores important mechanisms involved in islet cell hemostasis. Further clinical intervention strategies should be based in the concepts presented by the authors. However, better comprehension of the whole pathophysiological processes might be achieved with a schematic diagram.	1. addressed
5.	(Reviewed by 00073640): The subject of the review is very interesting and topical. The manuscript is very good structured and concise. However, to improve the manuscript the schematic representation (or table) of the involved molecules and their roles are highly recommended.	1. addressed

Comments to reviewers:

Many thanks to all reviewers for their overall positive comments and critical suggestions. The following has been incorporated within the manuscript ('32367-Revised word document') as revisions/corrections:

- 1) Reviewer 00504316: Schematic photography included
- 2) Reviewer 00070310: All articles within the reference section have now been listed with the PMID number and DOI.
- 3) Reviewer 00503542: Many thanks for the inputs and useful suggestions. All the suggested corrections have been considered and revised within the manuscript (yellow highlighted regions within the '32367-Revised word document').
- 4) Reviewer 02993037: Schematic diagram is included
- 5) Reviewer 00073640: Schematic representation has been included.