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Dr. Alisha DSouza Senior Review Editor JoVE

October 31, 2019

Dear Dr. DSouza.

Thank you for serving as the review editor for our manuscript entitled, *Generation of Mosaic Mammary Organoids by Differential Trypsinization*. We appreciated the time and attention that six reviewers spent in assessing our manuscript and uploaded you will find a point-by-point response to each comment. We also appreciated the editorial review and comments. Please find our point-by-point response below.

Best Regards,

Lindsay Hinck

#### **Editorial comments:**

Ludson Houce

You will find Editorial comments and Peer-Review comments listed below. Please read this entire email before making edits to your manuscript.

NOTE: Please include a line-by-line response to each of the editorial and reviewer comments in the form of a letter along with the resubmission.

## **Editorial Comments:**

- Please take this opportunity to thoroughly proofread the manuscript to ensure that there are no spelling or grammatical errors.
- Protocol Detail: Please note that your protocol will be used to generate the script for the video, and must contain everything that you would like shown in the video. Please add more specific details (e.g. button clicks for software actions, numerical values for settings, etc) to your protocol steps. There should be enough detail in each step to supplement the actions seen in the video so that viewers can easily replicate the protocol. Examples:
- 1) 1.1: Is the animal euthanized/anesthetized? Mention all surgical prep steps and anesthesia method. Mention all tools used and describe how sterility is maintained. What is the animal age, strain, sex?

We updated the first step of our protocol to include more details about how the animals are euthanized.

2) 1.1.: please clarify if the lymph nodes are to be included in the rest of the experiment. "Excise"/"extract" may be better words to use here.

We added detailed images to Figure 1 showing the lymph nodes.

3) 1.2,2.10: What is the composition of Digestion Media/ Maintenance Media?

All of the media recipes are listed in the excel speadsheet with recipes.

### • Protocol Highlight:

- 1) The highlighted steps should form a cohesive narrative, that is, there must be a logical flow from one highlighted step to the next.
- 2) Please highlight complete sentences (not parts of sentences). Include sub-headings and spaces when calculating the final highlighted length.
- 3) Notes cannot be filmed and should be excluded from highlighting.

Thank you, we made the changes to our highlighted selections.

• **Results:** Please add at least one paragraph of results text that explains your representative results in the context of the technique you describe; i.e. how do these results show the technique, suggestions about how to analyze the outcome etc. This text should be written in paragraph form under a "Representative Results" heading and should refer to all of the results figures. You may include the figure captions under this heading but the captions and figure text must be separate entities.

We included details about our results in the Representative Results section.

#### • Discussion:

1) JoVE articles are focused on the methods and the protocol, thus the discussion should be similarly focused. Please ensure that the discussion covers the following in detail and in paragraph form (3-6 paragraphs): 1) modifications and troubleshooting, 2) limitations of the technique, 3) significance with respect to existing methods, 4) future applications and 5) critical steps within the protocol.

2) Avoid numbered/bulleted lists in this section.

The discussion was rewritten to match the formatting requirements.

### • References:

1) Please make sure that your references comply with JoVE instructions for authors. Citation formatting should appear as follows: (For 6 authors or less list all authors. For more than 6 authors, list only the first author then *et al.*): [Lastname, F.I., LastName, F.I., LastName, F.I. Article Title. *Source*. **Volume** (Issue), FirstPage – LastPage, doi:DOI (YEAR).]
2) Please spell out journal names.

Yes, we changed the formatting.

- Commercial Language: JoVE is unable to publish manuscripts containing commercial sounding language, including trademark or registered trademark symbols (TM/R) and the mention of company brand names before an instrument or reagent. Examples of commercial sounding language in your manuscript are Matrigel,
- 1) Please use MS Word's find function (Ctrl+F), to locate and replace all commercial sounding language in your manuscript with generic names that are not company-specific. All commercial products should be sufficiently referenced in the table of materials/reagents. You may use the generic term followed by "(see table of materials)" to draw the readers' attention to specific commercial names.
- 2) Please check your figures as well.

Yes, we changed the use of Matrigel to ECM.

• Table of Materials: Please revise the table of the essential supplies, reagents, and equipment. The table

should include the name, company, and catalog number of all relevant materials/software in separate columns in an xls/xlsx file. Please include items such as animal strain, media used, etc.

We have made changes to make the tables more consistent.

• If your figures and tables are original and not published previously or you have already obtained figure permissions, please ignore this comment. If you are re-using figures from a previous publication, you must obtain explicit permission to re-use the figure from the previous publisher (this can be in the form of a letter from an editor or a link to the editorial policies that allows you to re-publish the figure). Please upload the text of the re-print permission (may be copied and pasted from an email/website) as a Word document to the Editorial Manager site in the "Supplemental files (as requested by JoVE)" section. Please also cite the figure appropriately in the figure legend, i.e. "This figure has been modified from [citation]."

We uploaded the Elsevier License that we obtained through RightsLink and cited Figure 2 as suggested above in the legend.