

October 6, 2016

Dr. Jaydev Upponi, Ph.D.
Science Editor, *Journal of Visualized Experiments*

Re: Manuscript 55346_R1_090216, "A method for orthotopic transplantation of lung cancer in mice"

Dear Dr. Jaydev Upponi,

Thank you for the invitation to resubmit the above manuscript. We appreciate the editorial comments to improve this manuscript. In the enclosed revised manuscript, I believe we have addressed the all of the editorial comments in regard to this manuscript. Our responses are below each comment and is italicized following ">>>".

Editorial Comments:

- Please keep the editorial comments from your previous revisions in mind as you revise your manuscript to address peer review comments. For instance, if formatting or other changes were made, commercial language was removed, etc., please maintain these overall manuscript changes.

>>> *Previous changes in response to editorial comments have been retained.*

- Animal use note: JoVE veterinary reviewers recommend using a scalpel, not scissors, to make initial incisions for survival procedures.

>>> *As noted in previous version, we have discussed this with our veterinarians at UTSW who noted that using scissors as described is safe. Due to the small incision that our procedure requires and the thinness of the skin, we feel that if safer for the mouse and operator if we pull up on the skin with forceps to elevate it away from chest wall and cut a small incision. Thus, this limits the risk of damage to the underlying chest wall and ribs.*

- The manuscript should receive copyediting for grammar and formatting issues. Some examples are listed here, although this is not a comprehensive list:

- Ranges of values, such as "3-5 min" or "5-10 mL" should be formatted as having a single hyphen and no spaces. Currently, the formatting varies from step to step (see, for example, steps 2.4 and 2.5).

>>> *The text has modified to the requested formatting changes.*

- SI units should be used consistently throughout (e.g. mL rather than ml, h rather than hrs, etc.).

>>> *The text has modified to the requested formatting changes.*

- Grammatical articles (e.g. a, an, the) are often missing throughout the Protocol section.

>>> *The text has modified to the requested formatting changes.*

-The Short Abstract should probably begin, "This protocol..." rather than "The protocol..."
Likewise, "This model" rather than "The model."

>>> *The text has modified to the requested formatting changes.*

-First sentence of Long Abstract: "Lung cancer is the most common cause of cancer death worldwide for which chemotherapy still remains the dominant mode of systemic therapy for most metastatic lung cancers."

>>> *We are not clear as to what needs to be modified in this comment.*

-Step 4.3: "Check the level of anesthesia by pinching the toes and monitor for withdrawal reflex."

>>> *We are not clear as to what needs to be modified in this comment.*

-Step 4.5: "Place anesthetized-mouse..." This hyphen is not necessary.

>>> *The text has modified to the requested formatting changes.*

-Step 5.1: "Place the mouse on the right lateral decubitus position" should be "in" rather than "on."

>>> *The text has modified to the requested formatting changes.*

-Step 5.12: The second sentence should be re-written in imperative tense.

>>> *The text has modified to the requested formatting changes.*

•Additional detail or clarification is needed:

-Step 4.5 instructs to use an electric clipper or trimmer to shave the fur of the mouse. Step 4.6 then instructs to "dispose of all materials used to shave the mouse." Do the authors really dispose of an electric trimmer after each shaving?

>>> *Sentence has been modified such that all "disposable" items be disposed.*

-Step 5.14 refers to steps 4.7-4.12, but section 4 ends with step 4.8.

>>> *The references in step 5.14 has been corrected.*

•If your figures and tables are original and not published previously, please ignore this comment. For figures and tables that have been published before, please include phrases such as "Re-print with permission from (reference#)" or "Modified from.." etc. And please send a copy of the re-print permission for JoVE's record keeping purposes.

>>> *All figures are original.*

•JoVE reference format requires that the DOIs are included, when available, for all references listed in the article. This is helpful for readers to locate the included references and obtain more information. Please note that often DOIs are not listed with PubMed abstracts and as such, may not be properly included when citing directly from PubMed. In these cases, please manually include DOIs in reference information.

>>> *We have been unable to find doi numbers for references #10, 11, and 19 including the actual articles. References 2, 7, and 8 are web pages and do not have doi numbers. All other references have doi numbers.*

•IMPORTANT: Please copy-edit the entire manuscript for any grammatical errors you may find. The text should be in American-English only. This editing should be performed by a native English speaker (or professional copyediting services) and is essential for clarity of the protocol and the manuscript. Please thoroughly review the language and grammar prior to resubmission. Your JoVE editor will not copy-edit your manuscript and any errors in your submitted revision may be present in the published version.

>>> *We have copy-edited our manuscript for any grammatical errors.*

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

>>> *Please see "CoverLetter_Response to Reviewers' Comments"*

Thank you again for the comments and the opportunity to resubmit our manuscript.

Sincerely,



James Kim, MD PhD