



**Mutation-agnostic RNA interference and engineered replacement rescues Tmc1-related hearing loss**

Yoichiro Iwasa, Miles Klimara, Hidekane Yoshimura, William Walls, Ryotaro Omichi, Cody West, Seiji Shibata, Paul Ranum and Richard Smith

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*Corresponding author(s): Prof. Richard JH Smith (University of Iowa)*

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**Review timeline:**

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Editorial Decision:	2022-11-09
Revision Received:	2022-12-09
Editorial Decision:	2022-12-12
Revision Received:	2022-12-14
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*Scientific Editor: Novella Guidi*

**Transaction Report:**

(Note: With the exception of the correction of typographical or spelling errors that could be a source of ambiguity, letters and reports are not edited. The original formatting of letters and referee reports may not be reflected in this compilation.)

No full Peer Review Process File is available with this article, as the authors have chosen not to make the review process public in this case.



Re: Life Science Alliance manuscript #LSA-2022-01592-T

Prof. Richard JH Smith  
University of Iowa  
Molecular Otolaryngology and Renal Research Laboratories, Department of  
Otolaryngology - Head and Neck Surgery  
Iowa City, IA 52242

Dear Dr. Smith,

Thank you for submitting your manuscript entitled "Mutation-agnostic RNA interference and engineered replacement rescues Tmc1-related hearing loss" to Life Science Alliance. The manuscript was assessed by expert reviewers, whose comments are appended to this letter. We invite you to submit a revised manuscript addressing the Reviewer comments.

To upload the revised version of your manuscript, please log in to your account:  
<https://lsa.msubmit.net/cgi-bin/main.plex>

You will be guided to complete the submission of your revised manuscript and to fill in all necessary information. Please get in touch in case you do not know or remember your login name.

While you are revising your manuscript, please also attend to the below editorial points to help expedite the publication of your manuscript. Please direct any editorial questions to the journal office.

The typical timeframe for revisions is three months. Please note that papers are generally considered through only one revision cycle, so strong support from the referees on the revised version is needed for acceptance.

When submitting the revision, please include a letter addressing the reviewers' comments point by point.

We hope that the comments below will prove constructive as your work progresses.

Thank you for this interesting contribution to Life Science Alliance. We are looking forward to receiving your revised manuscript.

Sincerely,

Novella Guidi, PhD  
Scientific Editor  
Life Science Alliance

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-- A letter addressing the reviewers' comments point by point.

-- An editable version of the final text (.DOC or .DOCX) is needed for copyediting (no PDFs).



-- High-resolution figure, supplementary figure and video files uploaded as individual files: See our detailed guidelines for preparing your production-ready images, <https://www.life-science-alliance.org/authors>

-- Summary blurb (enter in submission system): A short text summarizing in a single sentence the study (max. 200 characters including spaces). This text is used in conjunction with the titles of papers, hence should be informative and complementary to the title and running title. It should describe the context and significance of the findings for a general readership; it should be written in the present tense and refer to the work in the third person. Author names should not be mentioned.

-- By submitting a revision, you attest that you are aware of our payment policies found here: <https://www.life-science-alliance.org/copyright-license-fee>

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We encourage our authors to provide original source data, particularly uncropped/-processed electrophoretic blots and spreadsheets for the main figures of the manuscript. If you would like to add source data, we would welcome one PDF/Excel-file per figure for this information. These files will be linked online as supplementary "Source Data" files.

\*\*\*IMPORTANT: It is Life Science Alliance policy that if requested, original data images must be made available. Failure to provide original images upon request will result in unavoidable delays in publication. Please ensure that you have access to all original microscopy and blot data images before submitting your revision.\*\*\*

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Re: Life Science Alliance manuscript #LSA-2022-01592-TR

Prof. Richard JH Smith  
University of Iowa  
Molecular Otolaryngology and Renal Research Laboratories, Department of  
Otolaryngology - Head and Neck Surgery  
285 Newton Rd  
5270 Carver Biomedical Research Building  
Iowa City, IA 52242

Dear Dr. Smith,

Thank you for submitting your revised manuscript entitled "Mutation-agnostic RNA interference and engineered replacement rescues Tmc1-related hearing loss" to Life Science Alliance. The manuscript has been seen by the original reviewers whose comments are appended below. While the reviewers continue to be overall positive about the work in terms of its suitability for Life Science Alliance, some important issues remain.

Our general policy is that papers are considered through only one revision cycle; however, given that the suggested changes are relatively minor, we are open to one additional short round of revision. Please note that I will expect to make a final decision without additional reviewer input upon resubmission.

Please submit the final revision within one month, along with a letter that includes a point by point response to the remaining reviewer comments.

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Please pay attention to adhere to our editorial requirements for revisions:

#### A. THESE ITEMS ARE REQUIRED FOR REVISIONS

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#### B. MANUSCRIPT ORGANIZATION AND FORMATTING:

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Thank you for this interesting contribution to Life Science Alliance.

Sincerely,

Novella Guidi, PhD  
Scientific Editor  
Life Science Alliance

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RE: Life Science Alliance Manuscript #LSA-2022-01592-TRR

Prof. Richard JH Smith  
University of Iowa  
Molecular Otolaryngology and Renal Research Laboratories, Department of  
Otolaryngology - Head and Neck Surgery  
285 Newton Rd  
5270 Carver Biomedical Research Building  
Iowa City, IA 52242

Dear Dr. Smith,

Thank you for submitting your revised manuscript entitled "Mutation-agnostic RNA interference and engineered replacement rescues Tmc1-related hearing loss". We would be happy to publish your paper in Life Science Alliance pending final revisions necessary to meet our formatting guidelines.

Along with points mentioned below, please tend to the following:

- please use the [10 author names, et al.] format in your references (i.e. limit the author names to the first 10)
- please rename your EV figures as supplementary figures and update the figure callouts in the text as necessary; we do not have a limit on the number of supplementary figures
- please add your table legend to the main manuscript text
- please remove 'The Paper Explained' section

Figure Check:

-Figure 4 and Figure EV3, Figure EV4 look like there are duplicates throughout the figures; this is not allowed at LSA. Please remove duplicates and provide source data for these figures

If you are planning a press release on your work, please inform us immediately to allow informing our production team and scheduling a release date.

LSA now encourages authors to provide a 30-60 second video where the study is briefly explained. We will use these videos on social media to promote the published paper and the presenting author (for examples, see <https://twitter.com/LSAjournal/timelines/1437405065917124608>). Corresponding or first-authors are welcome to submit the video. Please submit only one video per manuscript. The video can be emailed to [contact@life-science-alliance.org](mailto:contact@life-science-alliance.org)

To upload the final version of your manuscript, please log in to your account:

<https://lsa.msubmit.net/cgi-bin/main.plex>

You will be guided to complete the submission of your revised manuscript and to fill in all necessary information. Please get in touch in case you do not know or remember your login name.

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**\*\*Submission of a paper that does not conform to Life Science Alliance guidelines will delay the acceptance of your manuscript.\*\***

**\*\*It is Life Science Alliance policy that if requested, original data images must be made available to the editors. Failure to provide original images upon request will result in unavoidable delays in publication. Please ensure that you have access to all original data images prior to final submission.\*\***

**\*\*The license to publish form must be signed before your manuscript can be sent to production. A link to the electronic license to publish form will be sent to the corresponding author only. Please take a moment to check your funder requirements.\*\***

**\*\*Reviews, decision letters, and point-by-point responses associated with peer-review at Life Science Alliance will be published online, alongside the manuscript. If you do want to opt out of having the reviewer reports and your point-by-point responses displayed, please let us know immediately.\*\***

Thank you for your attention to these final processing requirements. Please revise and format the manuscript and upload materials within 7 days.

Thank you for this interesting contribution, we look forward to publishing your paper in Life Science Alliance.

Sincerely,

Novella Guidi, PhD  
Scientific Editor  
Life Science Alliance

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4th Editorial Decision

14 December 2022

RE: Life Science Alliance Manuscript #LSA-2022-01592-TRRR

Prof. Richard JH Smith

University of Iowa  
Molecular Otolaryngology and Renal Research Laboratories, Department of  
Otolaryngology - Head and Neck Surgery  
285 Newton Rd  
5270 Carver Biomedical Research Building  
Iowa City, IA 52242

Dear Dr. Smith,

Thank you for submitting your Research Article entitled "Mutation-agnostic RNA interference and engineered replacement rescues Tmc1-related hearing loss". It is a pleasure to let you know that your manuscript is now accepted for publication in Life Science Alliance. Congratulations on this interesting work.

The final published version of your manuscript will be deposited by us to PubMed Central upon online publication.

Your manuscript will now progress through copyediting and proofing. It is journal policy that authors provide original data upon request.

Reviews, decision letters, and point-by-point responses associated with peer-review at Life Science Alliance will be published online, alongside the manuscript. If you do want to opt out of having the reviewer reports and your point-by-point responses displayed, please let us know immediately.

**\*\*\*IMPORTANT:** If you will be unreachable at any time, please provide us with the email address of an alternate author. Failure to respond to routine queries may lead to unavoidable delays in publication.\*\*\*

Scheduling details will be available from our production department. You will receive proofs shortly before the publication date. Only essential corrections can be made at the proof stage so if there are any minor final changes you wish to make to the manuscript, please let the journal office know now.

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Again, congratulations on a very nice paper. I hope you found the review process to be constructive and are pleased with how the manuscript was handled editorially. We look forward to future exciting submissions from your lab.

Sincerely,

Novella Guidi, PhD  
Scientific Editor  
Life Science Alliance