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Dear Arup Chakraborty and Reviewers,

Thank you for the careful and rapid review of our manuscript *“Mapping mutational effects along the evolutionary landscape of HIV envelope.”*

We were delighted to receive reviews that were both favorable and provided extremely helpful advice in terms of further strengthening the paper.

In response to those comments, we have elaborated on key questions in the text, added a few new panels to existing figures, and created some new figure supplements. Our changes are detailed in a point-by-point response to the reviewers. In addition, we have included a version of the manuscript that has changes tracked relative to the original submission.

Finally, in the methods, we now reference a GitHub repository with the computer code and LaTex files for the paper (<https://github.com/jbloomlab/EnvMutationalShiftsPaper>). We hope that this repository might be forked on the *eLife* GitHub, and have added text referring to such a fork similar to what we have been instructed to do for prior *eLife* papers.

Thanks for your time and consideration.

Sincerely,



Jesse D. Bloom, Ph.D.