

**From:** [Raffield, Laura](#)  
**To:** [Sedlazeck, Fritz](#)  
**Cc:** [jhspub](#); [April P. Carson](#); [Metcalf, Ginger A.](#); [TOPMed ACC Publications](#)  
**Subject:** RE: [EXTERNAL]RE: question re TOPMed structural variants manuscript...  
**Date:** Tuesday, January 17, 2023 4:28:58 PM

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**CAUTION: External Email \***

Thanks Fritz! Yes, totally makes sense that imputation server incorporation is not trivial at all. I was hoping allele frequency look-ups with Bravo might be easier near term, but I know nothing is easy for giant TOPMed projects .

Thanks for including me and JHS in this interesting work!

Laura

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**From:** Sedlazeck, Fritz <Fritz.Sedlazeck@bcm.edu>  
**Sent:** Tuesday, January 17, 2023 2:03 PM  
**To:** Raffield, Laura <laura\_raffield@unc.edu>  
**Cc:** Sedlazeck, Fritz <Fritz.Sedlazeck@bcm.edu>; [jhspub](#) <[jhspub@umc.edu](#)>; April P. Carson <[apcarson@umc.edu](#)>; Metcalf, Ginger A. <[metcalf@bcm.edu](#)>; TOPMed ACC Publications <[topmed-pubs@westat.com](#)>  
**Subject:** Re: [EXTERNAL]RE: question re TOPMed structural variants manuscript...

Thanks Laura,

I will work on this. @Bravo : not right now, but thats the next step. The imputation part is not trivial as you probably are aware of.

Thanks

Fritz

On Jan 17, 2023, at 11:00 AM, Raffield, Laura <[laura\\_raffield@unc.edu](#)> wrote:

Hi Fritz,

Great news re JHS approval already happening! I spoke with April and actually I will be the JHS representative for this one. I added myself to the spreadsheet. Thanks for updating those acknowledgements.

I looked through the paper and have no major concerns- a couple tiny comments below. Sounds like we are ready to submit from the JHS perspective, thanks for your patience.

1. The TOPLD manuscript is now out, please update that citation-<https://pubmed.ncbi.nlm.nih.gov/35504290/>

2. What is the kinship coefficient threshold for defining “founders” in KING? And how much does this reduce the total freeze 9 sample size limiting to founder populations?
3. The current manuscript doesn’t seem to include anything on the UK Biobank structural variation work- isn’t there already at least a preliminary callset from their WGS, as in<https://www.nature.com/articles/s41586-022-04965-x>?
4. I know you cite Taliun et al, but I think you need to give at least brief details on how genetic ancestry is defined in the methods here.
5. Are these structural variant calls going to be integrated into BRAVO so people can check the allele frequencies?

Best,  
Laura

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**From:** Sedlazeck, Fritz <[Fritz.Sedlazeck@bcm.edu](mailto:Fritz.Sedlazeck@bcm.edu)>  
**Sent:** Tuesday, January 17, 2023 9:52 AM  
**To:** jhspub <[jhspub@umc.edu](mailto:jhspub@umc.edu)>  
**Cc:** Sedlazeck, Fritz <[Fritz.Sedlazeck@bcm.edu](mailto:Fritz.Sedlazeck@bcm.edu)>; Raffield, Laura <[laura\\_raffield@unc.edu](mailto:laura_raffield@unc.edu)>; April P. Carson <[apcarson@umc.edu](mailto:apcarson@umc.edu)>; Metcalf, Ginger A. <[metcalf@bcm.edu](mailto:metcalf@bcm.edu)>; TOPMed ACC Publications <[topmed-pubs@westat.com](mailto:topmed-pubs@westat.com)>  
**Subject:** Re: [EXTERNAL]RE: question re TOPMed structural variants manuscript...

Ok updated.  
Thanks  
Fritz

On Jan 17, 2023, at 6:02 AM, jhspub <[jhspub@umc.edu](mailto:jhspub@umc.edu)> wrote:

Good morning,

Yes; please update the acknowledgment statement.

Best,  
Pam

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**From:** Sedlazeck, Fritz <[Fritz.Sedlazeck@bcm.edu](mailto:Fritz.Sedlazeck@bcm.edu)>  
**Sent:** Friday, January 13, 2023 5:04 PM  
**To:** jhspub <[jhspub@umc.edu](mailto:jhspub@umc.edu)>  
**Cc:** Raffield, Laura <[laura\\_raffield@unc.edu](mailto:laura_raffield@unc.edu)>; Sedlazeck, Fritz <[Fritz.Sedlazeck@bcm.edu](mailto:Fritz.Sedlazeck@bcm.edu)>; April P. Carson <[apcarson@umc.edu](mailto:apcarson@umc.edu)>; Metcalf, Ginger A. <[metcalf@bcm.edu](mailto:metcalf@bcm.edu)>; TOPMed ACC Publications

<[topmed-pubs@westat.com](mailto:topmed-pubs@westat.com)>

**Subject:** Re: [EXTERNAL]RE: question re TOPMed structural variants manuscript...

Dear Pam,

Thank you for this. Should we update the acknowledgement as indicated by Laura?

Thank you all for the fast turnaround and help with this.

Fritz

On Jan 13, 2023, at 3:38 PM, jhspub <[jhspub@umc.edu](mailto:jhspub@umc.edu)> wrote:

Good afternoon,

Thank you for your email. Our records indicate that this manuscript has already been submitted and reviewed. Please see the attached decision letter for your reference. Should you have any questions, please let us know.

Best,  
Pam

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**From:** Raffield, Laura <[laura\\_raffield@unc.edu](mailto:laura_raffield@unc.edu)>

**Sent:** Friday, January 13, 2023 9:59 AM

**To:** Sedlazeck, Fritz <[Fritz.Sedlazeck@bcm.edu](mailto:Fritz.Sedlazeck@bcm.edu)>; jhspub <[jhspub@umc.edu](mailto:jhspub@umc.edu)>

**Cc:** April P. Carson <[apcarson@umc.edu](mailto:apcarson@umc.edu)>; Metcalf, Ginger A. <[metcalf@bcm.edu](mailto:metcalf@bcm.edu)>; TOPMed ACC Publications <[topmed-pubs@westat.com](mailto:topmed-pubs@westat.com)>

**Subject:** [EXTERNAL]RE: question re TOPMed structural variants manuscript...

Hi Fritz,

Happy New Year! Hope you had a nice holiday break. Totally understand the difficulty of coordinating approvals across such a large # of studies. I think a reasonable solution would be to include JHS co-authors (will defer to April on who the two representatives should be), update JHS acknowledgments (current version below) and make sure

this is submitted for expedited review ASAP (those are done by email and should take less than a week, I think it would be ok for you to go ahead and submit before the TOPMed meeting Jan 26 if April agrees)... the jhspub team (cc'd) should be able to help (unless there is any record of a July review/submission?). Including the submitted paper files for April's reference, if I am one of the coauthors or the P&P reviewer I will be sure to take a more detailed read.

Thanks!  
Laura

The Jackson Heart Study (JHS) is supported and conducted in collaboration with Jackson State University (HHSN268201800013I), Tougaloo College (HHSN268201800014I), the Mississippi State Department of Health (HHSN268201800015I) and the University of Mississippi Medical Center (HHSN268201800010I, HHSN268201800011I and HHSN268201800012I) contracts from the National Heart, Lung, and Blood Institute (NHLBI) and the National Institute on Minority Health and Health Disparities (NIMHD). The authors also wish to thank the staffs and participants of the JHS.

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**From:** Sedlazeck, Fritz <[Fritz.Sedlazeck@bcm.edu](mailto:Fritz.Sedlazeck@bcm.edu)>  
**Sent:** Friday, January 13, 2023 9:14 AM  
**To:** Raffield, Laura <[laura\\_raffield@unc.edu](mailto:laura_raffield@unc.edu)>  
**Cc:** Sedlazeck, Fritz <[Fritz.Sedlazeck@bcm.edu](mailto:Fritz.Sedlazeck@bcm.edu)>; April P. Carson <[apcarson@umc.edu](mailto:apcarson@umc.edu)>; Metcalf, Ginger A. <[metcalf@bcm.edu](mailto:metcalf@bcm.edu)>; TOPMed ACC Publications <[topmed-pubs@westat.com](mailto:topmed-pubs@westat.com)>  
**Subject:** Re: question re TOPMed structural variants manuscript...

You don't often get email from [fritz.sedlazeck@bcm.edu](mailto:fritz.sedlazeck@bcm.edu). [Learn why this is important](#)

Dear Laura,  
Thank you for your email and sorry for the late reply. I took a longer holiday these season.

This manuscript is a larger collection of Topmed related Structural Variations and was compiled as part of the Topmed IRC. I see we reached out to jhspub in June 2022, but dont recall if we ever got a response. Sorry for this oversight , but as you can imagine this data set with 140000WGS includes many studies. In any case, how can we process this and resolve this in a timely manner? We hoped to be able to submit this manuscript before the Topmed meeting.

For the other studies we included two people as authors. Would this be you and Dr. Carson?  
Could you please add you both with information (affiliation, email and conflict of interest) at the end of this document: [https://docs.google.com/spreadsheets/d/1DcLd\\_1K\\_2704-ZzpL9wfbwkXuSw8NVHq7\\_PojlDnkYw/edit?usp=sharing](https://docs.google.com/spreadsheets/d/1DcLd_1K_2704-ZzpL9wfbwkXuSw8NVHq7_PojlDnkYw/edit?usp=sharing)

Thank you for your help  
Fritz

On Dec 23, 2022, at 1:16 PM, Raffield, Laura  
<[laura\\_raffield@unc.edu](mailto:laura_raffield@unc.edu)> wrote:

**\*\*\*CAUTION:\*\*\* This email is not from a BCM Source. Only click links or open attachments you know are safe.**

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Hi Fritz,

I noticed that you have submitted the TOPMed structural variant paper to the TOPMed website, but I do not believe this manuscript has been approved by JHS P&P (or at least no letter was attached). Has it been submitted and approved by 'jhspub' [jhspub@umc.edu](mailto:jhspub@umc.edu)? Also, the paper does not seem to include any JHS specific representatives, including from the coordinating center? Perhaps JHS PI Dr. Carson could be added and the paper could be submitted to P&P (before it is submitted to preprint server or journal)? Thanks for your

attention to this issue and for including JHS in  
this work.

Happy holidays!

Best,

Laura

Laura Raffield

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she / her / hers

<2021\_topmedSVPaper\_version

2\_Sedlazeck.pdf>

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