

Re: TOPMed Cohort: AA/CAC - P&P Approval

Taylor, Kent <ktaylor@lundquist.org>

Mon 8/16/2021 12:41 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

External Email - Use Caution

I approve the proposal

Kent

From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>**Sent:** Monday, August 16, 2021 8:05 AM**To:** Guo, Xiuqing <xguo@lundquist.org>; Li, Xiaohui <xiaohuili@lundquist.org>; Taylor, Kent <ktaylor@lundquist.org>**Cc:** Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>**Subject:** TOPMed Cohort: AA/CAC - P&P Approval

CAUTION - EXTERNAL EMAIL: Treat all links and attachments with extreme caution even if you recognize the sender.

Dear AA/CAC Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!

Margaret

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Please note that this e-mail is not secure (encrypted). If you do not wish to continue communication over unencrypted e-mail, please notify the sender of this message immediately. Continuing to send or respond to e-mail after receiving this message means you understand and accept this risk and wish to continue to communicate over unencrypted e-mail.

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penalties. Should you not be the intended recipient, please notify us by return email, and delete this message from your computer at once.

RE: TOPMed Cohort: Amish- P&P ApprovalMitchell, Braxton <bmitchel@som.umaryland.edu>

Fri 8/27/2021 2:35 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mg.harvard.edu>; Montasser, May <mmontass@som.umaryland.edu>; O'Connell, Jeff <joconnel@som.umaryland.edu>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

External Email - Use Caution

Hi Margaret,,
This is approved for Amish.
Brackie

From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mg.harvard.edu>

Sent: Friday, August 27, 2021 11:46 AM

To: Mitchell, Braxton <bmitchel@som.umaryland.edu>; Montasser, May <mmontass@som.umaryland.edu>; O'Connell, Jeff <joconnel@som.umaryland.edu>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: Re: TOPMed Cohort: Amish- P&P Approval

Thanks Dr.Braxton

Do you require any specific detail from our end to obtain the P&P approval?

Thanks
Margaret

From: Mitchell, Braxton <bmitchel@som.umaryland.edu>

Sent: Monday, August 16, 2021 11:53 AM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mg.harvard.edu>; Montasser, May <mmontass@som.umaryland.edu>; O'Connell, Jeff <joconnel@som.umaryland.edu>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: RE: TOPMed Cohort: Amish- P&P Approval

External Email - Use Caution

Will confirm shortly.
Brackie

From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mg.harvard.edu>

Sent: Monday, August 16, 2021 11:34 AM

To: Mitchell, Braxton <bmitchel@som.umaryland.edu>; Montasser, May <mmontass@som.umaryland.edu>; O'Connell, Jeff <joconnel@som.umaryland.edu>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: TOPMed Cohort: Amish- P&P Approval

CAUTION: This message originated from a non UMB, UMSOM, FPI, or UMMS email system. Whether the sender is known or not known, hover over any links before clicking and use caution opening attachments.

Dear Amish Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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**Research
with Heart.**

Memorandum

To: Paul S. de Vries
From: Richey Sharrett, for the ARIC Publications Committee
CC: ARIC Publications Committee; ARIC Coordinating Center
Subject: Review of ARIC MS # T9318
Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Date: Monday, August 23, 2021

On behalf of the ARIC Steering Committee, I am happy to inform you that the above-referenced paper has been approved.

ARIC Publication Policies Require:

1. To comply with NIH Public Access Policy, all ARIC manuscripts are required to be submitted to PubMed Central upon acceptance for publication. If your paper is accepted for publication in a journal that deposits all final published articles in PubMed Central, no author involvement is required. If the journal does not automatically deposit all final manuscripts into PubMed Central, you (or a co-author) **MUST** deposit the final peer-reviewed manuscript in PubMed Central via the NIH Manuscript Submission System (NIHMS): <http://www.nihms.nih.gov/>. More information regarding NIH Public Access Policy can be found here: <http://publicaccess.nih.gov/>
2. Enclosed are the instructions to authors, per our policy, as well as a sentence acknowledging ARIC staff, which should be added to the paper.
3. Please remember to add reference to "ARIC," in either the title of the paper or the abstract, so that the study can be identified by reference searches. In addition, when the paper has been published, please email the citation and pdf to both Jordan Darden (aricpub@unc.edu) and Kendra Hay (aricjhu@jhu.edu).
4. Please follow procedures local to your center for archiving a copy of the programs which created this paper. At Johns Hopkins provide a copy to Natalie Daya (ndaya1@jhu.edu).

Congratulations and good luck with this interesting paper's submission.

Decision: **Approved**

Re: TOPMed Cohort: BioMe - P&P Approval

Loos, Ruth <ruth.loos@mssm.edu>

Mon 8/16/2021 2:54 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Wang, Zhe <zhe.wang@mssm.edu>; Preuss, Michael <michael.preuss@mssm.edu>**Cc:** Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>**External Email - Use Caution**

Hello Margaret,

We do not have a P&P, but I approve.

Thanks for including BioMe.

All the best,
Ruth

From: "Selvaraj, Margaret Sunitha" <MSSELVARAJ@mgh.harvard.edu>**Date:** Monday, August 16, 2021 at 5:36 PM**To:** "Loos, Ruth" <ruth.loos@mssm.edu>, "Wang, Zhe" <zhe.wang@mssm.edu>, "Preuss, Michael" <michael.preuss@mssm.edu>**Cc:** "Pradeep Natarajan (pradeep@broadinstitute.org)" <pradeep@broadinstitute.org>, "Peloso, Gina Marie" <gpeloso@bu.edu>, "S. Margaret Sunitha" <sunitha@broadinstitute.org>**Subject:** TOPMed Cohort: BioMe - P&P ApprovalUSE CAUTION: External
Message.

Dear BioMe Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
MargaretThe information in this e-mail is intended only for the person to whom it is addressed. If you believe this e-mail was sent to you in error and the e-mail contains patient information, please contact the Mass General Brigham Compliance HelpLine at <http://www.massgeneralbrigham.org/complianceline> . If the e-mail was sent to you in error but does not contain patient information, please contact the sender and properly dispose of the e-mail.

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respond to e-mail after receiving this message means you understand and accept this risk and wish to continue to communicate over unencrypted e-mail.

Re: TOPMed Cohort: CARDIA - P&P Approval

Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

Mon 8/16/2021 12:27 PM

To: Fornage, Myriam <Myriam.Fornage@uth.tmc.edu>; jwilkins@nm.org <jwilkins@nm.org>
Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Dear Dr.Myriam

I have included the suggested changes to the manuscript. Thank you.

Best,
Margaret

From: Fornage, Myriam <Myriam.Fornage@uth.tmc.edu>

Sent: Monday, August 16, 2021 12:11 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; jwilkins@nm.org <jwilkins@nm.org>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: RE: TOPMed Cohort: CARDIA - P&P Approval

External Email - Use Caution

Dear Margaret:

Very nice work. Attached are some minor edits. Please make sure to include CARDIA acknowledgments (attached).

Once it is done, you can consider the manuscript approved for CARDIA.

Best wishes for the submission,
Myriam

From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

Sent: Monday, August 16, 2021 10:39 AM

To: Fornage, Myriam <Myriam.Fornage@uth.tmc.edu>; jwilkins@nm.org

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: TOPMed Cohort: CARDIA - P&P Approval

****** EXTERNAL EMAIL ******

Dear CARDIA Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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Please note that this e-mail is not secure (encrypted). If you do not wish to continue communication over unencrypted e-mail, please notify the sender of this message immediately. Continuing to send or respond to e-mail after receiving this message means you understand and accept this risk and wish to continue to communicate over unencrypted e-mail.

RE: TOPMed Cohort: CFS - P&P Approval

Redline, Susan,M.D.,M.P.H. <SREDLINE@BWH.HARVARD.EDU>

Mon 8/16/2021 5:28 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Cade, Brian,PH.D. <bcade@bwh.harvard.edu>
Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Thanks again for your wonderful paper- it is beautifully written. Other than two small typos, I had nothing to add. So, as PI of CFS I approve-

Good luck-

From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

Sent: Monday, August 16, 2021 11:40 AM

To: Redline, Susan,M.D.,M.P.H. <SREDLINE@BWH.HARVARD.EDU>; Cade, Brian,PH.D. <bcade@bwh.harvard.edu>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: TOPMed Cohort: CFS - P&P Approval

Dear CFS Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

Re: Record update from chs-nhlbi.org

Natarajan, Pradeep, M.D. <PNATARAJAN@mgh.harvard.edu>

Mon 8/2/2021 8:07 PM

To: Joshua C. Bis <joshbis@uw.edu>

Cc: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Peloso, Gina Marie <gpeloso@bu.edu>

Thanks!

From: Joshua C. Bis <joshbis@uw.edu>

Date: Monday, August 2, 2021 at 6:38 PM

To: Natarajan, Pradeep, M.D. <PNATARAJAN@mgh.harvard.edu>

Subject: Fwd: Record update from chs-nhlbi.org

External Email - Use Caution

See below for confirmation that CHS approved the lipids paper.

. josh

Begin forwarded message:

From: chsnhlbi@u.washington.edu

Date: August 2, 2021 at 3:19:36 PM PDT

To: joshbis@uw.edu

Subject: Record update from chs-nhlbi.org

Reply-To: chsnhlbi@u.washington.edu

Hello, authors.

Congratulations! Your penultimate draft, Whole genome sequence analysis of blood lipid levels in >66,000 individuals, was approved by the Steering Committee. Please see the updated status and details regarding your next steps at the following link:
<https://chs-nhlbi.org/node/8798>.

Please let me know if you have any questions.

Best,
Ellen

Ellen Terry
Publications Coordinator
CHS-NHLBI.org
chsnhlbi@u.washington.edu

NOTES: Please make sure you are logged on to the CHS website before clicking the above link. This message was sent to the first author and all CHS co-authors.

<https://chs-nhlbi.org/node/8798>.

RE: TOPMed Cohort: DHS - P&P Approval

Barry Freedman <bfreedma@wakehealth.edu>

Tue 8/17/2021 7:41 AM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Donald W. Bowden <dbowden@wakehealth.edu>; Nicholette (Palmer) Allred <nallred@wakehealth.edu>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>; Barry Freedman <bfreedma@wakehealth.edu>

External Email - Use Caution

Hi Margaret,

Thanks – I don't see that Nichole replied and Don is out of office. The DHS approves submission – you are good from our end.

Appreciate all of the hard work. Thanks, Barry

Barry I. Freedman, MD, FACP

John H. Felts III Professor
Chief, Section on Nephrology
Wake Forest School of Medicine
Medical Center Boulevard
Winston-Salem, NC 27157-1053
p 336.716.6192 / f 336.716.4318
bfreedma@wakehealth.edu

Chief Medical Officer
Health Systems Management, Inc.
For Wake Forest Outpatient Dialysis
605 Cotton Street, Suite 200
Winston-Salem, NC 27101
For Emory Outpatient Dialysis
610 Northside Drive
Atlanta, GA 30318
p 336.748.0575 / f 336.722.9759

From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

Sent: Monday, August 16, 2021 11:44 AM

To: Donald W. Bowden <dbowden@wakehealth.edu>; Nicholette (Palmer) Allred <nallred@wakehealth.edu>; Barry Freedman <bfreedma@wakehealth.edu>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: [EXTERNAL] TOPMed Cohort: DHS - P&P Approval

Dear DHS Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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Please note that this e-mail is not secure (encrypted). If you do not wish to continue communication over unencrypted e-mail, please notify the sender of this message immediately. Continuing to send or respond to e-mail after receiving this message means you understand and accept this risk and wish to continue to communicate over unencrypted e-mail.



S. Margaret Sunitha <sunitha@broadinstitute.org>

FW: TOPMed Lipids 66K freeze 8 paper for review

Peloso, Gina Marie <gpeloso@bu.edu>
To: "S. Margaret Sunitha" <sunitha@broadinstitute.org>

Tue, Aug 3, 2021 at 2:07 PM

From: Ramachandran, Vasan
Sent: Tuesday, July 27, 2021 1:32 PM
To: Peloso, Gina Marie <gpeloso@bu.edu>; Cupples, L Adrienne <adrienne@bu.edu>
Subject: RE: TOPMed Lipids 66K freeze 8 paper for review

Nope.

Just make sure the TOPMed acknowledgments are correct for FHS (consistent with our other FHS TOPMed papers).

Thanks, Gina.

Best,

Vasan

From: Peloso, Gina Marie <gpeloso@bu.edu>
Sent: Tuesday, July 27, 2021 1:29 PM
To: Ramachandran, Vasan <vasan@bu.edu>; Cupples, L Adrienne <adrienne@bu.edu>
Subject: RE: TOPMed Lipids 66K freeze 8 paper for review

Sounds good. I'll add in Yuxuan as well.

Is there anything else I need to do for FHS approval?

Thank you,

Gina

From: Ramachandran, Vasan
Sent: Tuesday, July 27, 2021 1:25 PM
To: Peloso, Gina Marie <gpeloso@bu.edu>; Cupples, L Adrienne <adrienne@bu.edu>
Subject: RE: TOPMed Lipids 66K freeze 8 paper for review

Thanks, Gina.

I had listed/included (in the online google sheet) Nancy Heard-Costa and myself as additional people, who made this project feasible.

Sincerely,

Vasan

From: Peloso, Gina Marie <gpeloso@bu.edu>
Sent: Tuesday, July 27, 2021 1:23 PM
To: Ramachandran, Vasan <vasan@bu.edu>; Cupples, L Adrienne <adrienne@bu.edu>
Subject: FW: TOPMed Lipids 66K freeze 8 paper for review

Dear Vasan and Adrienne,

I would like to nominate the following authors from FHS for this manuscript:

Gina Peloso

Yuxuan Wang (my PhD student working on this project)

L. Adrienne Cupples

Please let me know if this is ok or there should be any further authors on this TOPMed paper and I will update the authorship list.

Thank you,

[Quoted text hidden]

Re: TOPMed Lipids 66K freeze 8 paper for review

Lisa Yanek <lryanek@jhmi.edu>

Mon 8/9/2021 10:17 AM

To: Natarajan, Pradeep, M.D. <PNATARAJAN@mgh.harvard.edu>**Cc:** Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Peloso, Gina Marie <gpeloso@bu.edu>**External Email - Use Caution**

Hi all,

The manuscript and figures look great! This is now approved by our GeneSTAR steering committee.

Best,
Lisa

From: Natarajan, Pradeep, M.D. <PNATARAJAN@mgh.harvard.edu>**Sent:** Tuesday, July 27, 2021 2:49 PM**To:** Lisa Yanek <lryanek@jhmi.edu>**Cc:** Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Peloso, Gina Marie <gpeloso@bu.edu>**Subject:** FW: TOPMed Lipids 66K freeze 8 paper for review**External Email - Use Caution**

From: Natarajan, Pradeep, M.D. <PNATARAJAN@mgh.harvard.edu>**Date:** Sunday, July 25, 2021 at 2:40 PM**To:** xihaoli@g.harvard.edu <xihaoli@g.harvard.edu>, Li, Zilin <li@hsph.harvard.edu>, pampana.akhil@gmail.com <pampana.akhil@gmail.com>, david.zhang@penntmedicine.upenn.edu <david.zhang@penntmedicine.upenn.edu>, joseph.park@penntmedicine.upenn.edu <joseph.park@penntmedicine.upenn.edu>, Rader, Daniel <rader@penntmedicine.upenn.edu>, cclaurie@uw.edu <cclaurie@uw.edu>, goncalo@umich.edu <goncalo@umich.edu>, Eric.Boerwinkle@uth.tmc.edu <Eric.Boerwinkle@uth.tmc.edu>, bmitchel@som.umaryland.edu <bmitchel@som.umaryland.edu>, ruth.loos@mssm.edu <ruth.loos@mssm.edu>, Myriam.Fornage@uth.tmc.edu <Myriam.Fornage@uth.tmc.edu>, Redline, Susan, M.D., M.P.H. <SREDLINE@BWH.HARVARD.EDU>, psaty@uw.edu <psaty@uw.edu>, dbowden@wakehealth.edu <dbowden@wakehealth.edu>, vasan@bu.edu <vasan@bu.edu>, skardia@umich.edu <skardia@umich.edu>, stella7@gmail.com <stella7@gmail.com>, jhe@tulane.edu <jhe@tulane.edu>, rmathias@jhmi.edu <rmathias@jhmi.edu>, Robert.Kaplan@einstein.yu.edu <Robert.Kaplan@einstein.yu.edu>, donna.arnett@uky.edu <donna.arnett@uky.edu>, acorrea@umc.edu <acorrea@umc.edu>, jrotter@labiomed.org <jrotter@labiomed.org>, Ellinor, Patrick Thomas, M.D., Ph.D. <ellinor@mgh.harvard.edu>, john.blanger@utrgv.edu <john.blanger@utrgv.edu>, stephen_mcgarvey@brown.edu <stephen_mcgarvey@brown.edu>, rao@wustl.edu <rao@wustl.edu>, clk@fredhutch.org <clk@fredhutch.org>, ssr4n@virginia.edu <ssr4n@virginia.edu>, cristen@umich.edu <cristen@umich.edu>, Xihong Lin <xlin@hsph.harvard.edu>**Cc:** Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>, Peloso, Gina Marie

<gpeloso@bu.edu>

Subject: TOPMed Lipids 66K freeze 8 paper for review

Dear colleagues,

We're excited to share the TOPMed Lipids 66K freeze 8 paper draft for your review/feedback.

Manuscript:

https://www.dropbox.com/s/781llxflu02p9jm/Lipid66K_Freeze8_manuscript_V6_GMP_PN.docx?dl=0

Supp Text: https://www.dropbox.com/s/drvyhp00kaf1p0i/Supplementary_Final.docx?dl=0

Supp Tables: https://www.dropbox.com/s/qmrj4ckamfth3fm/SupplementaryTables_Final.xlsx?dl=0

Please provide feedback by **Mon (Aug 9)**. Please also send to your corresponding P&Ps (or let us know if you'd like to do this & which email address to send it to). Please also note that we need you to identify authors for your respective authorship group (3 per group as we have done previously with our working group), so please fill out

<https://docs.google.com/spreadsheets/d/1wOvVV3B63fZ663Y2fR7H6ecbGpv2WI6V8oQua7xdkPY/edit#gid=0>

If we do not hear back from you by Mon (Aug 9), we assume that you have no additional feedback, no additional P&P is necessary for your cohort, and your authorship group is finalized.

Please do not hesitate to reach out with any questions/comments. Many thanks for your collaboration and we look forward to your constructive feedback (as well as the administrative guidance – P&Ps and authorship)!

Margaret, Gina, and Pradeep

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Re: TOPMed Cohort: GENOA - P&P Approval

Jennifer Smith <smjenn@umich.edu>

Mon 8/16/2021 11:58 AM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>**Cc:** skardia@umich.edu <skardia@umich.edu>; zhaowei@umich.edu <zhaowei@umich.edu>; Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>**External Email - Use Caution**

Hi Margaret,

We don't have a formal P&P for GENOA. However, I will review the manuscript that you sent out today and let you know if GENOA needs any changes.

Thank you!

Jen

~~~~~  
Jennifer A. Smith, PhD, MPH

Associate Professor, Department of Epidemiology, School of Public Health

Research Associate Professor, Survey Research Center, Institute for Social Research

University of Michigan

1415 Washington Heights, Room 2631

Ann Arbor, MI 48109-2029

Tel: (734) 615-9455

Fax: (734) 764-3192

Email: [smjenn@umich.edu](mailto:smjenn@umich.edu)  
~~~~~

On Mon, Aug 16, 2021 at 11:45 AM Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu> wrote:

Dear GENOA Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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does not contain patient information, please contact the sender and properly dispose of the e-mail.

Please note that this e-mail is not secure (encrypted). If you do not wish to continue communication over unencrypted e-mail, please notify the sender of this message immediately. Continuing to send or respond to e-mail after receiving this message means you understand and accept this risk and wish to continue to communicate over unencrypted e-mail.

Re: TOPMed Cohort: GenSalt - P&P Approval

Pradeep Natarajan <pradeep@broadinstitute.org>

Thu 8/19/2021 10:06 AM

To: He, Jiang <jhe@tulane.edu>**Cc:** Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Kelly, Tanika N <tkelly@tulane.edu>; Sun, Xiao <xsun10@tulane.edu>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>**External Email - Use Caution**

Thanks for clarifying.

Pradeep



On Aug 19, 2021, at 8:45 AM, He, Jiang <jhe@tulane.edu> wrote:

We do not have. Thanks,

Jiang

From: Pradeep Natarajan <pradeep@broadinstitute.org>**Sent:** Thursday, August 19, 2021 6:19 AM**To:** He, Jiang <jhe@tulane.edu>**Cc:** Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Kelly, Tanika N <tkelly@tulane.edu>; Sun, Xiao <xsun10@tulane.edu>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>**Subject:** Re: TOPMed Cohort: GenSalt - P&P Approval

External Sender. Be aware of links, attachments and requests.

Some cohorts (not all) have a separate committee to formally approve papers and proposals on behalf of the cohort. Does GenSalt have such a committee?

Thanks

Pradeep

On Aug 18, 2021, at 10:44 PM, He, Jiang <jhe@tulane.edu> wrote:

Tell us exact what you need. Thanks,

Jiang

From: Pradeep Natarajan <pradeep@broadinstitute.org>**Sent:** Wednesday, August 18, 2021 8:06 PM**To:** Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; He, Jiang <jhe@tulane.edu>; Kelly, Tanika N <tkelly@tulane.edu>; Sun, Xiao <xsun10@tulane.edu>

Cc: Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: Re: TOPMed Cohort: GenSalt - P&P Approval

External Sender. Be aware of links, attachments and requests.

Hi folks,

Please let us know if GenSalt requires a separate P&P process (in addition to TOPMed's usual P&P process).

Thanks
Pradeep

From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>
Date: Wednesday, August 18, 2021 at 10:31 AM
To: jhe@tulane.edu <jhe@tulane.edu>, tkelly@tulane.edu <tkelly@tulane.edu>, xsun10@tulane.edu <xsun10@tulane.edu>
Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>, Peloso, Gina Marie <gpeloso@bu.edu>, S. Margaret Sunitha <sunitha@broadinstitute.org>
Subject: Re: TOPMed Cohort: GenSalt - P&P Approval

Dear GenSalt Team

Do you require any specific information from our end to move forward with the P&P approval process?

Please let us know, so that I can send you the details.

Thanks,
Margaret

From: Selvaraj, Margaret Sunitha
Sent: Monday, August 16, 2021 11:49 AM
To: jhe@tulane.edu <jhe@tulane.edu>; tkelly@tulane.edu <tkelly@tulane.edu>; xsun10@tulane.edu <xsun10@tulane.edu>
Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>
Subject: TOPMed Cohort: GenSalt - P&P Approval

Dear GenSalt Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9

Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals

Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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RE: TOPMed Cohort: GOLDN - P&P Approval

Arnett, Donna K. <donna.arnett@uky.edu>

Mon 8/16/2021 1:48 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

Cc: Claas, Steve <Steven.Claas@uky.edu>; Claas, Steve <Steven.Claas@uky.edu>

External Email - Use Caution

Approved.

From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

Sent: Monday, August 16, 2021 11:47 AM

To: stella7@gmail.com; Arnett, Donna K. <donna.arnett@uky.edu>; berthahidalgo@uab.edu

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: TOPMed Cohort: GOLDN - P&P Approval

CAUTION: External Sender

Dear GOLDN Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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S. Margaret Sunitha <sunitha@broadinstitute.org>

HCHS - Coauthors addition - Whole genome sequence analysis of blood lipid levels in >66,000 individuals

S. Margaret Sunitha <sunitha@broadinstitute.org>

Mon, Aug 16, 2021 at 12:41 PM

To: Robert Kaplan <robert.kaplan@einsteinmed.org>

Cc: "North, Kari E" <kari_north@unc.edu>, "Graff, Misa" <migraff@email.unc.edu>

Thanks for your approval Dr. Kaplan. We are composing a response draft for the comments, and will send you the revision soon we compile them together.

Best,
Margaret

On Mon, Aug 16, 2021 at 12:12 PM Robert Kaplan <robert.kaplan@einsteinmed.org> wrote:

Dear Margaret, please see below, approval by Kari and me serves as sign-off for the SOL study.

Do you have a revised version of the MS?

Thank you

Robert

Robert Kaplan, PhD
Albert Einstein College of Medicine
718-430-4076 (New York)
206-667-6913 (Seattle)

From: Robert Kaplan <robert.kaplan@einsteinmed.org>

Sent: Wednesday, August 4, 2021 2:37 PM

To: S. Margaret Sunitha <sunitha@broadinstitute.org>; North, Kari E <kari_north@unc.edu>

Cc: Pradeep Natarajan <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Graff, Misa <migraff@email.unc.edu>

Subject: Re: HCHS - Coauthors addition - Whole genome sequence analysis of blood lipid levels in >66,000 individuals

Dear Margaret,

Here are comments on the draft manuscript. Overall looks good. As per our SOL policy, review by Kari and me serves as the "cohort" approval.

Good luck, thanks for the invitation to be a coauthor.

Robert

1, I would recommend to specify in the text of the MS how many of the "findings" of the study were actually just repeating what was already known from GWAS. For example this statement below, it does not really explain the real story which was that if you look at the number of novel variants, the number was tiny and we discovered very little in this WGS analysis with regard to the search for novel variants. I think the paper should point this out, it seems like this important fact is obscured somewhat.

"Among genome-wide significant variants, we tabulated 'known-position' (variant previously 122associated), 'known-loci' (variants not previously significantly associated with the corresponding lipid phenotype but within 123500 kb of a knownlocus, thereby representing additional allelic heterogeneity), and 'novel' variants(variants not in a 124known lipid locus) (SupplementaryTable 3) (line 121-4).

2, I do not understand this point about confounding. What does the size of the study have to do with the likelihood of confounding? *" However, whether residual confoundingexistsbecause of the relatively limited sample size of TOPMed compared to vastly larger lipid GWAS, increasingly in non-Europeans, is not known (line 283-4)"*

3, Fig. 2 Summary of single variant genome wide association: Please clarify the legend to explain what is included in the inner rings (our findings from TOPMed). Did this set of variants include only those meeting GW significance, or also the "suggestive" levels of significance?

4, Use of lipid treatments is important to understand better, perhaps this should be added to ST1. Somewhere I recommend that we should show the frequency of lipid treatments in each cohort. Maybe it was included somewhere but I missed it.

5, About half of the WGS data in SOL that is being used here was funded by NHGRI through the CCDG program, so that should be included in the supplemental text funding acknowledgment.

Robert Kaplan, PhD
Albert Einstein College of Medicine
718-430-4076 (New York)
206-667-6913 (Seattle)

From: S. Margaret Sunitha <sunitha@broadinstitute.org>
Sent: Wednesday, August 4, 2021 12:13 PM
To: North, Kari E <kari_north@unc.edu>
Cc: Robert Kaplan <robert.kaplan@einsteinmed.org>; Pradeep Natarajan <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Graff, Misa <migraff@email.unc.edu>
Subject: Re: HCHS - Coauthors addition - Whole genome sequence analysis of blood lipid levels in >66,000 individuals

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Dear Dr. Robert Kaplan
Thanks for your reply, I am resending the email that was earlier circulated. You will be able to see the manuscript main and supplementary materials in the dropbox link.

Best,
Margaret

Dear colleagues,

We're excited to share the TOPMed Lipids 66K freeze 8 paper draft for your review/feedback.
Manuscript: https://www.dropbox.com/s/781llxflu02p9jm/Lipid66K_Freeze8_manuscript_V6_GMP_PN.docx?dl=0

Supp Text: https://www.dropbox.com/s/drvyhp00kaf1p0i/Supplementary_Final.docx?dl=0

Supp Tables: https://www.dropbox.com/s/qmrj4ckamfth3fm/SupplementaryTables_Final.xlsx?dl=0

Please provide feedback by **Mon (Aug 9)**. Please also send to your corresponding P&Ps (or let us know if you'd like to do this & which email address to send it to). Please also note that we need you to identify authors for your respective authorship group (3 per group as we have done previously with our working group), so please fill out <https://docs.google.com/spreadsheets/d/1wOvVV3B63fZ663Y2fR7H6ecbGpv2Wl6V8oQua7xdkPY/edit#gid=0>

If we do not hear back from you by **Mon (Aug 9)**, we assume that you have no additional feedback, no additional P&P is necessary for your cohort, and your authorship group is finalized.

Please do not hesitate to reach out with any questions/comments. Many thanks for your collaboration and we look forward to your constructive feedback (as well as the administrative guidance – P&Ps and authorship)!

Margaret, Gina, and Pradeep

On Wed, Aug 4, 2021 at 3:08 PM North, Kari E <kari_north@unc.edu> wrote:

I do not recall seeing the paper.

If already circulated, please forward to me and Misa for review.

Kari

Kari E. North
Professor of Epidemiology
Director, CVD Genetic Epidemiology Computational Laboratory
Gillings School of Global Public Health
University of North Carolina
123 W. Franklin Street, Building C, Suite 4217
CB#8050
Chapel Hill, NC 27599-8050
919-966-2148; Fax 919-966-9800
<http://hlbgenepi.web.unc.edu>

--

From: Robert Kaplan <robert.kaplan@einsteinmed.org>

Date: Wednesday, August 4, 2021 at 2:59 PM

To: "S. Margaret Sunitha" <sunitha@broadinstitute.org>, Kari North <kari_north@unc.edu>

Cc: Pradeep Natarajan <pradeep@broadinstitute.org>, "gpeloso@bu.edu" <gpeloso@bu.edu>, "Selvaraj, Margaret Sunitha" <MSSELVARAJ@mgh.harvard.edu>, Mariaelisa Graff <migraff@email.unc.edu>

Subject: Re: HCHS - Coauthors addition - Whole genome sequence analysis of blood lipid levels in >66,000 individuals

Dear Margaret,

Thanks for contacting me again. I have not been able to access the manuscript. Seems like the website required me to request some kind of permission, and then nothing happened after that. Could you please just email me the text of the paper, at least I can take a look at that.

Thanks,
Robert

Robert Kaplan, PhD

Albert Einstein College of Medicine
718-430-4076 (New York)
206-667-6913 (Seattle)

From: S. Margaret Sunitha <sunitha@broadinstitute.org>
Sent: Wednesday, August 4, 2021 11:20 AM
To: North, Kari E <kari_north@unc.edu>
Cc: Robert Kaplan <robert.kaplan@einsteinmed.org>; Pradeep Natarajan <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Graff, Misa <migraff@email.unc.edu>
Subject: Re: HCHS - Coauthors addition - Whole genome sequence analysis of blood lipid levels in >66,000 individuals

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Thank you.
Margaret

On Wed, Aug 4, 2021 at 2:15 PM North, Kari E <kari_north@unc.edu> wrote:

Done.

Kari E. North
Professor of Epidemiology
Director, CVD Genetic Epidemiology Computational Laboratory
Gillings School of Global Public Health
University of North Carolina
123 W. Franklin Street, Building C, Suite 4217
CB#8050
Chapel Hill, NC 27599-8050
919-966-2148; Fax 919-966-9800
<http://hlbgenepi.web.unc.edu>
--

From: "S. Margaret Sunitha" <sunitha@broadinstitute.org>
Date: Wednesday, August 4, 2021 at 2:06 PM
To: Kari North <kari_north@unc.edu>
Cc: "robert.kaplan@einsteinmed.org" <robert.kaplan@einsteinmed.org>, Pradeep Natarajan <pradeep@broadinstitute.org>, "gpeloso@bu.edu" <gpeloso@bu.edu>, "Selvaraj, Margaret Sunitha" <MSSELVARAJ@mgh.harvard.edu>, Mariaelisa Graff <migraff@email.unc.edu>
Subject: Re: HCHS - Coauthors addition - Whole genome sequence analysis of blood lipid levels in >66,000 individuals

That's great to hear from you, can you please fill the google form, I shared the file with you and you can make edits to it.

Thank you,
Margaret

On Wed, Aug 4, 2021 at 2:01 PM North, Kari E <kari_north@unc.edu> wrote:

Very sorry about this! I'm happy to participate as an author and also Mariaelisa Graff, copies here. Thanks! Kari

Sent from my iPhone

On Aug 4, 2021, at 1:47 PM, S. Margaret Sunitha <sunitha@broadinstitute.org> wrote:

Looping in Kari North.

Dear Kari

It would be great to hear back from the HCHS team about the list of co-authors to be included for our manuscript. We need to have the full authors list to obtain approvals from each cohort.

Therefore, obtaining the co-authors from your team ASAP will be helpful to move forward.

Thanks!

Margaret, Pradeep and Gina

On Fri, Jul 30, 2021 at 1:06 PM S. Margaret Sunitha <sunitha@broadinstitute.org> wrote:

Dear Dr.Robert Kaplan

We recently circulated co-authorship entry google documents for the manuscript titled as "Whole genome sequence analysis of blood lipid levels in >66,000 individuals". We would like to let you know that the document can be accessed from the following link. It would be great if you could add the co-authors names for this paper.

We are hoping you can address the author additions by Mon (8/2) as it should only take a few moments.

<https://docs.google.com/spreadsheets/d/1wOvVV3B63fZ663Y2fR7H6ecbGpv2Wl6V8oQua7xdkPY/edit#gid=0>

To begin cohort P&P reviews, most require the full authorship model. There will be sufficient time to provide feedback in parallel with the cohort P&P reviews.

Thanks!

Margaret, Pradeep and Gina

RE: TOPMed Cohort: HyperGEN - P&P Approval

Arnett, Donna K. <donna.arnett@uky.edu>

Mon 8/16/2021 1:14 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Irvin, Marguerite Ryan <irvinr@uab.edu>
Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

External Email - Use Caution

HyperGEN Approves.

From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>
Sent: Monday, August 16, 2021 1:00 PM
To: Arnett, Donna K. <donna.arnett@uky.edu>; Irvin, Marguerite Ryan <irvinr@uab.edu>
Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>
Subject: TOPMed Cohort: HyperGEN - P&P Approval

CAUTION: External Sender

Dear HyperGEN Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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Please note that this e-mail is not secure (encrypted). If you do not wish to continue communication over unencrypted e-mail, please notify the sender of this message immediately. Continuing to send or respond to e-mail after receiving this message means you understand and accept this risk and wish to continue to communicate over unencrypted e-mail.

RE: TOPMed Cohort: MGH_AF - P&P Approval

Ellinor, Patrick Thomas, M.D., Ph.D. <ellinor@mgh.harvard.edu>

Mon 8/16/2021 2:57 PM

To: Pradeep Natarajan <pradeep@broadinstitute.org>; Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>
Cc: lubitz@broadinstitute.org <lubit@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Great work and thanks for including us.

No P&P to worry about at MGH, thanks,
Patrick

From: Pradeep Natarajan <pradeep@broadinstitute.org>

Sent: Monday, August 16, 2021 2:16 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

Cc: Ellinor, Patrick Thomas, M.D., Ph.D. <ellinor@mgh.harvard.edu>; lubitz@broadinstitute.org; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: Re: TOPMed Cohort: MGH_AF - P&P Approval

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I assume no additional P&P approval for MGH-AF right?
Sorry, going through the requisite TOPMed administrative steps.

Pradeep



On Aug 16, 2021, at 1:02 PM, Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu> wrote:

Dear MGH-AF Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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RE: TOPMed Cohort: SAFS - P&P Approval

Joanne Curran <joanne.curran@utrgv.edu>

Mon 8/16/2021 1:10 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; John Blangero <john.blangero@utrgv.edu>; Ravindranath Duggirala <ravindranath.duggirala@utrgv.edu>**Cc:** Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>**External Email - Use Caution**

Hi Margaret,

There is no P&P approval required for SAFS.

Thanks,
Joanne**Joanne E. Curran, Ph.D.**

Professor

Lacks Valley Stores Ltd. Endowed Professor

Department of Human Genetics

South Texas Diabetes and Obesity Institute

1 West University Blvd,

Brownsville, TX 78520

+1 (956) 882-7532

joanne.curran@utrgv.edu

utrgv.edu

uthealthrgv.org

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From: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>**Sent:** Monday, August 16, 2021 12:07 PM**To:** John Blangero <john.blangero@utrgv.edu>; Joanne Curran <joanne.curran@utrgv.edu>; Ravindranath Duggirala <ravindranath.duggirala@utrgv.edu>**Cc:** Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>**Subject:** TOPMed Cohort: SAFS - P&P Approval**External Mail**

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Dear SAFS Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9
Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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Re: TOPMed Cohort: Samoan - P&P Approval

Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

Mon 8/16/2021 2:30 PM

To: McGarvey, Stephen <stephen_mcgarvey@brown.edu>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Dear Dr.Stephen,

We have made those changes to the manuscript supplementary table.

Thank you,
Margaret

From: McGarvey, Stephen <stephen_mcgarvey@brown.edu>

Sent: Monday, August 16, 2021 2:28 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>

Cc: Pradeep Natarajan (pradeep@broadinstitute.org) <pradeep@broadinstitute.org>; Peloso, Gina Marie <gpeloso@bu.edu>; S. Margaret Sunitha <sunitha@broadinstitute.org>

Subject: Re: TOPMed Cohort: Samoan - P&P Approval

External Email - Use Caution

Dear Margaret,

Subject to the changes I sent to Gina Peloso, we approve the manuscript for submission. To remind you our requested change was the following:

we request you use the word 'Samoan' in Suppl Table instead of the acronym 'SAS'.

best regards,

Steve McGarvey

Stephen T. McGarvey, PhD, MPH

Professor of Epidemiology and of Anthropology

International Health Institute

Brown University School of Public Health

Fellow, American Association for the Advancement of Science (AAAS)

121 S. Main St, Room 258

email: Stephen_McGarvey@brown.edu

On Mon, Aug 16, 2021 at 1:07 PM Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu> wrote:

Dear Samoan Team

We are in the process of obtaining P&P approvals from each cohort for the manuscript that we circulated recently.

TopMed Proposal Title: Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9

Proposal Id: 9318

Manuscript Title: Whole genome sequence analysis of blood lipid levels in >66,000 individuals
Target journal : Nature Genetics

It would be great to hear from you on the status of the P&P approval.

Thanks!
Margaret

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TEL +886 4 2359 2525 FAX +886 4 2359 5046 Website <http://www.vghtc.gov.tw>

April 21, 2020

Margaret Sunitha Selvaraj

Associate Professor

Broad Institute of Harvard and MIT

sunitha@broadinstitute.org

Subject: TOPMed THRV Collaboration Letter 9318

Dear Dr. Selvaraj,

We are pleased to hear about your plans to utilize data from the THRV Study to conduct your proposed project 9318 entitled: *Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9*.

We are happy to share THRV phenotype and whole genome sequence data with you and other TOPMed working group investigators working on this project and to collaborate on the analysis and interpretation of results of this study. We understand that results generated from this project, using data from THRV and other participating studies, will be reported and disseminated in peer-reviewed journals and at national conferences and that we will be informed in advance of each intended use. We also understand that we will have the opportunity to approve or disapprove the specific use cases. You agreed to 5 THRV Authors, those Authors are: Yii-Der Ida Chen, Lisa de la Fuentes, Chii-Min Hwa, Lee-Ming Chuang, and Ren-Hua Chung. Please acknowledge receipt of this letter via e-mail to Ms. Brigidann Cooper at brigidann.cooper@lundquist.org.

Sincerely,

Wayne Sheu (SAPPHIRE & THRV PI)

D.C. Rao (THRV PI)

Ida Chen (THRV Co-PI)





TOPMed

Published on *NHLBI Trans-Omics for Precision Medicine* (<https://www.nhlbiwgs.org>)

Whole genome sequence analysis of plasma lipids in TOPMed Freeze 9

Submitted by sunitha@broadin... on Tue, 04/07/2020 - 9:12am

Proposal id:	9318
Submitted:	Tue, 04/07/2020 - 9:12am
Proposed by:	Selvaraj, Margaret Sunitha
Proposer's email address:	sunitha@broadinstitute.org
Please describe the aims of the proposal:	<p>The aim of the proposal is to analyze the whole genome sequencing (WGS) data from TOPMed Freeze9. Specific objectives include</p> <ul style="list-style-type: none">Use of WGS data as a tool to identify novel loci associated with the phenotype of interest (blood lipid traits).To test the association of novel common and rare variants with blood lipid profile.To test the association of rare variants in both coding and non-coding regions with blood lipid profile more in detail.To test the association of rare variants separated by ethnic groups.To test the association of rare variants separated by gender.Fine mapping of significantly associated variants using eQTL and cis-regulatory element datasetsValidating the associated variants with external WGS cohort dataset
Proposal co-authors (Last, First):	Li, Zilin Li, Xihao Peloso, Gina Lin, Xihong Natarajan, Pradeep Correa, Adolfo Wilson, James Lange, Leslie

Mitchell, Braxton D
 Perry, James
 O'Connell, Jeff
 Psaty, Bruce
 Tracy, Russell
 Kooperberg, Charles
 Reiner, Alex
 Rao, D.C
 Chen, Yii-Der Ida
 Saleheen, Danish
 Arnett, Donna K
 Ellinor, Patrick
 Lubitz, Steven
 Boerwinkle, Eric
 Alanna C. Morrison
 Rotter, Jerome
 Rich, Stephen
 He, Jiang
 Hixson, James
 Blangero, John
 Curran, Joanne
 Mathias, Rasika
 Yanek, Lisa
 Kaplan, Robert
 North, Kari
 Loos, Ruth J.F
 Kenny, Eimear
 McGarvey, Stephen
 Weeks, Daniel E.
 Redline, Susan
 Ramachandran , Vasan S
 Cupples, L. Adrienne
 Fornage, Myriam
 Lloyd-Jones, Donald
 Arnett, Donna K
 Stella Aslibekyan

Is this a “single PI” proposal?:	No
List phenotypes/traits:	Plasma levels of total cholesterol; low-density lipoprotein cholesterol; high-density lipoprotein cholesterol; triglycerides; non- high-density lipoprotein cholesterol
Will your proposed analyses include all genomes sequenced through the overall TOPMed program, or a subset?:	Subset of genomes

If subset, please specify:	Only genomes with the phenotypes of interest.
Brief description of project analysis plans or lookup requested (½-1 ½ page):	<p>Background:</p> <p>There are more than 250 loci been associated with plasma lipid levels, interestingly common variants identified by GWAS explain only 10-15% of trait variance for lipids. The Common Disease-Rare Variant hypothesis proposes that the combined effect of a number of low frequency variants with large effect sizes accounts for some of the missing heritability. Previous studies show that a detail exploration at the common/rare variants in both coding and non-coding regions from larger cohorts could uncover novel findings associated with lipid traits. Moreover most of large scale GWAS studies have been carried out primarily on white followed by black populations and certain individual studies have focused on asian and hispanic population using both whole genome and exom sequencing data. This shows the need for a large scale multi-ethnic study which would help us understand the importance of association between novel variants and clinically recognized lipid traits.</p> <p>Analysis Plan:</p> <p>In the proposed project we aim to study the TOPMed freeze9 WGS dataset of autosomes with the goal of uncovering novel loci associated with lipid traits. In comparison with the previous TOPMed freeze datasets, the current data stands out on the number of samples from different ancestries and this gives us the scope to identify novel loci. In order to achieve this aim, we propose to study the data using multiple variant groupings. For the complete analysis we would use a sample set for which the lipid traits were quantified. The analysis of variants will be grouped into single variant analysis and rare variant analysis based on minor allele frequency (MAF).</p> <p>We will first harmonize the phenotypes of interest (Lipid traits - TC, TG, LDL, HDL) by merging datasets from different cohorts based on sample ids. Duplicate samples and samples less than 18 years of age will be removed. The raw lipid variables will be adjusted based on statin values and the fasting status. Next we will inverse rank normalization of the residuals of each race within each cohort scaled by the standard deviation of the trait adjusted for (Covariates) age, age2, sex, PC1-11, and recruitment site, where appropriate. Additional adjustment for founder mutation in APOB, APOC3 for certain cohort will be carried out.</p> <p>The Genetic relationship matrix (GRM) with PCA and relatedness (IBD) estimates will be downloaded from TOPMed common repository. The sparse kinship matrix sub-stetted to samples with lipid traits will be used in the following association analysis. Additionally annotations from VEP, LOFTEE, FAVOR and WGSa will be used for aggregating variants into groups.</p> <p>Single variant analysis will be carried out with variants of minor allele count (MAC) greater than 20. Variants belong to the class of common (MAF\geq5%), low-frequency (1%</p>

How may this analysis overlap other TOPMed projects:	This project is primarily focused on the analysis of freeze9 data from TOPMed, primarily on lipid traits. The project will add additional information on novel loci that are associated. The project does not overlap with any of the other studies. We will collaborate with the investigators of the lipids working group on analysis and publications.
What are your current plans for publication and authorship:	The findings will be published in peer-reviewed journals. We will include investigators who contribute data for these analyses, as well as investigators from the lipids working group who contribute to the design, conduct, analysis, interpretation, and/or writing of the manuscript as authors. We will publish under the banner of the TOPMed Lipids Working Group.
Will data be shared across institutions? Explain.:	Data will only be shared through standard TOPMed protocols to TOPMed investigators through the Exchange Area. Only summary statistics will be shared outside of TOPMed investigators.
Will individual-level data be distributed to investigators outside TOPMed? Explain.:	No
Primary Working Group:	Lipids [1]
Choose all other relevant working groups:	Lipids [1]
TOPMed Projects associated with this proposal:	AA_CAC [2] AFGen [3] Amish [4] BioMe [5] CARDIA [6] CFS [7] CHS [8] FHS [9] GeneSTAR [10] GenSalt [11] GOLDN [12] HCHS_SOL [13] HyperGEN_GENOA [14] JHS [15] MESA [16] PROMIS [17] SAFS [18] Samoan [19] THRIV [20] VTE [21]

[WHI](#) [22]

PI Contact:	I have contacted the PIs of the studies selected above after obtaining Working Group approval (or waiver) and indicated such to those PI. Each PI has either (a) agreed to a collaboration involving use of their study's data in the proposed paper or (b) did not respond to the request within two weeks. Please note that the proposal should be discussed and approved by a Working Group before contacting the study PI for their permission to use their study. I will also select specific consent groups from each study, using the 'Request data sets' form, as they become available.
Date actually posted:	Tuesday, April 7, 2020 - 9:12am
Scientific review status:	Pending review
Data Request Status:	In Progress
Overall Status:	In Progress

Source URL (modified on 04/07/2020 - 9:11am):

<https://www.nhlbiwgs.org/paperproposals/9318-whole-genome-sequence-analysis-plasma-lipids-topmed-freeze-9>

Links

- [1] <https://www.nhlbiwgs.org/working-group/54>
- [2] https://www.nhlbiwgs.org/group/AA_CAC#AA_CAC
- [3] <https://www.nhlbiwgs.org/group/AFGen#AFGen>
- [4] <https://www.nhlbiwgs.org/group/Amish#Amish>
- [5] <https://www.nhlbiwgs.org/group/BioMe#BioMe>
- [6] <https://www.nhlbiwgs.org/group/CARDIA#CARDIA>
- [7] <https://www.nhlbiwgs.org/group/CFS#CFS>
- [8] <https://www.nhlbiwgs.org/group/CHS#CHS>
- [9] <https://www.nhlbiwgs.org/group/FHS#FHS>
- [10] <https://www.nhlbiwgs.org/group/GeneSTAR#GeneSTAR>
- [11] <https://www.nhlbiwgs.org/group/GenSalt#GenSalt>
- [12] <https://www.nhlbiwgs.org/group/GOLDN#GOLDN>
- [13] https://www.nhlbiwgs.org/group/HCHS_SOL#HCHS_SOL
- [14] https://www.nhlbiwgs.org/group/HyperGEN_GENOA#HyperGEN_GENOA
- [15] <https://www.nhlbiwgs.org/group/JHS#JHS>
- [16] <https://www.nhlbiwgs.org/group/MESA#MESA>
- [17] <https://www.nhlbiwgs.org/group/PROMIS#PROMIS>
- [18] <https://www.nhlbiwgs.org/group/SAFS#SAFS>
- [19] <https://www.nhlbiwgs.org/group/Samoan#Samoan>
- [20] <https://www.nhlbiwgs.org/group/THRV#THRV>
- [21] <https://www.nhlbiwgs.org/group/VTE#VTE>
- [22] <https://www.nhlbiwgs.org/group/WHI#WHI>

FW: Ms4205 memo

Pradeep Natarajan <pradeep@broadinstitute.org>

Sat 8/14/2021 10:12 PM

To: Selvaraj, Margaret Sunitha <MSSELVARAJ@mgh.harvard.edu>; Peloso, Gina Marie <gpeloso@bu.edu> 1 attachments (197 KB)

Ms4205 memo.doc;

External Email - Use Caution

From: P&P Committee <p&p@WHI.org>**Date:** Friday, August 13, 2021 at 4:23 PM**To:** 'pradeep@broadinstitute.org' <pradeep@broadinstitute.org>, Charles Kooperberg <clk@fredhutch.org>**Cc:** P&P Committee <p&p@WHI.org>**Subject:** Ms4205 memo

Dear Charles and Pradeep,

Please see the attached memo regarding Ms4205.

Best,
Lindsey**Lindsey Bull**

Women's Health Initiative – Clinical Coordinating Center

Fred Hutch, Seattle WA

P&P@whi.org