Notice of Award



RESEARCH Department of Health and Human Services National Institutes of Health NIH

NATIONAL INSTITUTE OF BIOMEDICAL IMAGING AND BIOENGINEERING

Grant Number: 5R01EB015611-06 FAIN: R01EB015611

Principal Investigator(s): PETER V KOCHUNOV, PHD

Project Title: Solar-Eclipse Computational Tools for Imaging Genetics

Paffrath, Dennis Joseph AVP, Sponsored Programs Administration University of Maryland, Baltimore 620 West Lexington Street, 4129 Baltimore, MD 212011508

Award e-mailed to: nga@ordmail.umaryland.edu

Period Of Performance: Budget Period: 08/01/2018 – 07/31/2019 Project Period: 08/01/2012 – 07/31/2020

Dear Business Official:

The National Institutes of Health hereby awards a grant in the amount of \$400,001 (see "Award Calculation" in Section I and "Terms and Conditions" in Section III) to UNIVERSITY OF MARYLAND BALTIMORE in support of the above referenced project. This award is pursuant to the authority of 42 USC 241 42 CFR 52 and is subject to the requirements of this statute and regulation and of other referenced, incorporated or attached terms and conditions.

Acceptance of this award including the "Terms and Conditions" is acknowledged by the grantee when funds are drawn down or otherwise obtained from the grant payment system.

Each publication, press release, or other document about research supported by an NIH award must include an acknowledgment of NIH award support and a disclaimer such as "Research reported in this publication was supported by the National Institute Of Biomedical Imaging And Bioengineering of the National Institutes of Health under Award Number R01EB015611. The content is solely the responsibility of the authors and does not necessarily represent the official views of the National Institutes of Health." Prior to issuing a press release concerning the outcome of this research, please notify the NIH awarding IC in advance to allow for coordination.

Award recipients must promote objectivity in research by establishing standards that provide a reasonable expectation that the design, conduct and reporting of research funded under NIH awards will be free from bias resulting from an Investigator's Financial Conflict of Interest (FCOI), in accordance with the 2011 revised regulation at 42 CFR Part 50 Subpart F. The Institution shall submit all FCOI reports to the NIH through the eRA Commons FCOI Module. The regulation does not apply to Phase I Small Business Innovative Research (SBIR) and Small Business Technology Transfer (STTR) awards. Consult the NIH website

<u>http://grants.nih.gov/grants/policy/coi/</u> for a link to the regulation and additional important information.

If you have any questions about this award, please contact the individual(s) referenced in Section IV.

Sincerely yours,

Florence Turska Grants Management Officer NATIONAL INSTITUTE OF BIOMEDICAL IMAGING AND BIOENGINEERING

Additional information follows

SECTION I – AWARD DATA – 5R01EB015611-06	SECTION I - /	AWARD D	ATA - 5R01	EB015611-06
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Award Calculation (U.S. Dollars)	
Salaries and Wages	\$116,623
Fringe Benefits	\$33,407
Personnel Costs (Subtotal)	\$150,030
Consultant Services	\$7,000
Materials & Supplies	\$3,500
Travel	\$3,500
Other	\$5,336
Subawards/Consortium/Contractual Costs	\$138,330
Federal Direct Costs	\$307,696
Federal F&A Costs	\$92,305
Approved Budget	\$400,001
Total Amount of Federal Funds Obligated (Federal Share)	\$400,001
TOTAL FEDERAL AWARD AMOUNT	\$400,001
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AMOUNT OF THIS ACTION (FEDERAL SHARE)

\$400,001

	SUMMARY TOTALS FOR ALL YEARS		
YR THIS AWARD CUMULATIVE TOTALS		CUMULATIVE TOTALS	
6	\$400,001	\$400,001	
7	\$400,001	\$400,001	

Recommended future year total cost support, subject to the availability of funds and satisfactory progress of the project

Fiscal Information:

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IC	CAN	2018	2019
EB	8015183	\$400,001	\$400,001

Recommended future year total cost support, subject to the availability of funds and satisfactory progress of the project

NIH Administrative Data: PCC: HBIG / OC: 414E / Released: Were Name 08/08/2018 07:02:00 PM

SECTION II - PAYMENT/HOTLINE INFORMATION - 5R01EB015611-06

For payment and HHS Office of Inspector General Hotline information, see the NIH Home Page at http://grants.nih.gov/grants/policy/awardconditions.htm

SECTION III – TERMS AND CONDITIONS – 5R01EB015611-06

This award is based on the application submitted to, and as approved by, NIH on the above-titled project and is subject to the terms and conditions incorporated either directly or by reference in the following:

- a. The grant program legislation and program regulation cited in this Notice of Award.
- b. Conditions on activities and expenditure of funds in other statutory requirements, such as those included in appropriations acts.
- c. 45 CFR Part 75.

- d. National Policy Requirements and all other requirements described in the NIH Grants Policy Statement, including addenda in effect as of the beginning date of the budget period.
- e. Federal Award Performance Goals: As required by the periodic report in the RPPR or in the final progress report when applicable.
- f. This award notice, INCLUDING THE TERMS AND CONDITIONS CITED BELOW.

(See NIH Home Page at http://grants.nih.gov/grants/policy/awardconditions.htm for certain references cited above.)

Research and Development (R&D): All awards issued by the National Institutes of Health (NIH) meet the definition of "Research and Development" at 45 CFR Part§ 75.2. As such, auditees should identify NIH awards as part of the R&D cluster on the Schedule of Expenditures of Federal Awards (SEFA). The auditor should test NIH awards for compliance as instructed in Part V, Clusters of Programs. NIH recognizes that some awards may have another classification for purposes of indirect costs. The auditor is not required to report the disconnect (i.e., the award is classified as R&D for Federal Audit Requirement purposes but non-research for indirect cost rate purposes), unless the auditee is charging indirect costs at a rate other than the rate(s) specified in the award document(s).

An unobligated balance may be carried over into the next budget period without Grants Management Officer prior approval.

This grant is subject to Streamlined Noncompeting Award Procedures (SNAP).

This award is subject to the requirements of 2 CFR Part 25 for institutions to receive a Dun & Bradstreet Universal Numbering System (DUNS) number and maintain an active registration in the System for Award Management (SAM). Should a consortium/subaward be issued under this award, a DUNS requirement must be included. See

<u>http://grants.nih.gov/grants/policy/awardconditions.htm</u> for the full NIH award term implementing this requirement and other additional information.

This award has been assigned the Federal Award Identification Number (FAIN) R01EB015611. Recipients must document the assigned FAIN on each consortium/subaward issued under this award.

Based on the project period start date of this project, this award is likely subject to the Transparency Act subaward and executive compensation reporting requirement of 2 CFR Part 170. There are conditions that may exclude this award; see http://grants.nih.gov/grants/policy/awardconditions.htm for additional award applicability information.

In accordance with P.L. 110-161, compliance with the NIH Public Access Policy is now mandatory. For more information, see NOT-OD-08-033 and the Public Access website: <u>http://publicaccess.nih.gov/</u>.

In accordance with the regulatory requirements provided at 45 CFR 75.113 and Appendix XII to 45 CFR Part 75, recipients that have currently active Federal grants, cooperative agreements, and procurement contracts with cumulative total value greater than \$10,000,000 must report and maintain information in the System for Award Management (SAM) about civil, criminal, and administrative proceedings in connection with the award or performance of a Federal award that reached final disposition within the most recent five-year period. The recipient must also make semiannual disclosures regarding such proceedings. Proceedings information will be made publicly available in the designated integrity and performance system (currently the Federal Awardee Performance and Integrity Information System (FAPIIS)). Full reporting requirements and procedures are found in Appendix XII to 45 CFR Part 75. This term does not apply to NIH fellowships.

Treatment of Program Income:

SECTION IV - EB Special Terms and Conditions - 5R01EB015611-06

Clinical Trial Indicator: No

This award does not support any NIH-defined Clinical Trials. See the NIH Grants Policy Statement Section 1.2 for NIH definition of Clinical Trial.

CONSORTIUM/CONTRACTUAL COSTS

This award includes funds for consortium activity with:

- University of Texas Health Science Center at San Antonio
- University of Oxford, United Kingdom

Consortia are to be established and administered as described in the NIHGPS section 15 Consortium Agreements. http://grants.nih.gov/grants/policy/nihgps/HTML5/section 15/15 consortium agreements.htm

SALARY CAP

None of the funds in this award shall be used to pay the salary of an individual at a rate in excess of the current salary cap per year. Therefore, this award and/or future years are adjusted accordingly, if applicable. Current salary cap levels can be found at the following URL's:<u>https://grants.nih.gov/grants/guide/notice-files/NOT-OD-18-137.html</u>

GRADUATE STUDENT COMPENSATION

The maximum amount NIH will award for compensation of a graduate student (salary, fringe benefits and tuition remission) receiving support from a research grant is the zero-level Kirschstein-NRSA stipend in effect when NIH issues the grant award (see current levels posted at https://grants.nih.gov/grants/guide/notice-files/NOT-OD-18-175.html

The NIBIB home page is http://www.nibib.nih.gov/

STAFF CONTACTS

The Grants Management Specialist is responsible for the negotiation, award and administration of this project and for interpretation of Grants Administration policies and provisions. The Program Official is responsible for the scientific, programmatic and technical aspects of this project. These individuals work together in overall project administration. Prior approval requests (signed by an Authorized Organizational Representative) should be submitted in writing to the Grants Management Specialist. Requests may be made via e-mail.

Grants Management Specialist: Angelos Bacas Email: ab329b@nih.gov Phone: (301) 451-4785 Fax: (301) 451-5735

Program Official: Grace Peng Email: penggr@mail.nih.gov Phone: 301-451-4778 Fax: 301-480-1614

SPREADSHEET SUMMARY GRANT NUMBER: 5R01EB015611-06

INSTITUTION: UNIVERSITY OF MARYLAND BALTIMORE

Budget	Year 6	Year 7
Salaries and Wages	\$116,623	\$116,623
Fringe Benefits	\$33,407	\$33,407
Personnel Costs (Subtotal)	\$150,030	\$150,030
Consultant Services	\$7,000	\$7,000
Materials & Supplies	\$3,500	\$3,500
Travel	\$3,500	\$3,500

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Other	\$5,336	\$5,336
Subawards/Consortium/Contractual Costs	\$138,330	\$138,330
TOTAL FEDERAL DC	\$307,696	\$307,696
TOTAL FEDERAL F&A	\$92,305	\$92,305
TOTAL COST	\$400,001	\$400,001

Facilities and Administrative Costs	Year 6	Year 7
F&A Cost Rate 1	54.5%	54.5%
F&A Cost Base 1	\$169,367	\$169,367
F&A Costs 1	\$92,305	\$92,305

Project Title: Solar-Eclipse Computational Tools for Imaging Ge	enetics
Grant Number: 5R01EB015611-06 Project/Grant Period: 08/01/2012 - 07/31/2020	
Reporting Period: 08/01/2017 - 07/31/2018	Requested Budget Period: 08/01/2018 - 07/31/2019
Report Term Frequency: Annual	Date Submitted: 05/17/2018
Program Director/Principal Investigator Information:	Recipient Organization:
PETER V KOCHUNOV , MS MS PHD Phone number: (410) 402-6110 Email: Personal Info	UNIVERSITY OF MARYLAND BALTIMORE UNIVERSITY OF MARYLAND BALTIMORE 620 W LEXINGTON ST, 4TH FL BALTIMORE, MD 212011508 DUNS: 188435911 EIN: 1526002036A1 RECIPIENT ID:
Change of Contact PD/PI: N/A	
Administrative Official:	Signing Official:
MARIE COOLAHAN 620 W. Lexington Street 4th. Floor Baltimore, MD 21201 Phone number: 410-706-0011 Email: m_coolahan@umaryland.edu	MARIE COOLAHAN 620 W. Lexington Street 4th. Floor Baltimore, MD 21201 Phone number: 410-706-0011 Email: m_coolahan@umaryland.edu
Human Subjects: No	Vertebrate Animals: No
hESC: No	Inventions/Patents: No

B.1 WHAT ARE THE MAJOR GOALS OF THE PROJECT?

Aim 1. To shift imaging genetics toward richer, higher resolution imaging analyses and denser genome examination through high performance computing. The shift in imaging genetics toward WGS genotyping and high-resolution multimodal imaging data necessitates improvement of computational efficiency of likelihood calculations. To speed calculations up by 105-6-fold we will implement high-performance computing: novel data decompositions and single-step algorithmic techniques and integration of Massive Universal Linear Model (MULM) GPU library of genetic tools for massively parallel genetic analyses. These developments will enable interactive WGS and GWAS analyses for Big Data project by implementing SOLAR-Eclipse in web-analyses portals such as HCP Dashboard and ENIGMAVis.

Aim 2. To accelerate data sharing and replication in imaging genetics. Demands for high-performance computing and greater reproducibility and transparency in scientific research require a new data format optimized for imaging genetics applications and easy sharing of provenance. Expanding on the work of the International Neuroinformatics Coordinating Facility's Neuroimaging Data Sharing Task Force, we have assembled a panel of community experts to develop a draft of imaging genetic format based on existing neuroimaging formats along with extending on-going provenance efforts to imaging genetic research. We will submit this format and API for formal registration with NIF.

Aim 3. To implement and exploit empirical kinship methods. Empirical kinship algorithms that directly measure the degree of shared genetic variance (such as those used in GCTA/ REACTA and MEGHA) will be integrated for performing polygenic and Quantitative Trait Loci Linkage (QTL-L) analyses in the related, unrelated and mega-genetic samples. We propose to re-invent the Quantitative Trait Loci Linkage (QTL-L) methods for localizing QTLs based on simple empirical similarity in larger (1cM) regions of DNA instead of per-locus GWA-SNP analysis. This will answer questions such as localization of chromosomal segments that are responsible for normal and disorder-related variability in neuroimaging traits. Empirical QTL-L analyses will power chromosomal localization studies with no sharing of raw genotypes for our Big Data partners.

Aim 4. Interactive improvement of developed tools in collaboration with 'big data' partners. Methods developed in Aims 1-3 will 'push' the scientific aims of our Big Data partners, who committed a large (N=10K) sample and effort for three collaborative studies. The high-performance imaging genetics computing and empirical QTL-L techniques (Aims 1 and 3) will be honed by performing the largest genetic localization analyses with ENIGMA, ACP and GOBS projects, and by integrating voxel-wise GWAS analyses in web-analysis portals. Likewise, the utility of new format for data and workflow sharing (Aim 2) will be honed for multi-site Big Data research. Finally, we will develop and rank pioneering resting-state FMRI endophenotypes for Big Data research by demonstrating consistent heritability across samples and pleiotropy with mental disorders. The feedback gathered from our partners will sharpen SOLAR-Eclipse tools for imaging genetics community. We will continue to develop annual workshop at Imaging Genetics Conference to educate our users and disseminate new methods.

B.1.a Have the major goals changed since the initial competing award or previous report?

No

B.2 WHAT WAS ACCOMPLISHED UNDER THESE GOALS?

File uploaded: GoalsAchieved.pdf

B.3 COMPETITIVE REVISIONS/ADMINISTRATIVE SUPPLEMENTS

For this reporting period, is there one or more Revision/Supplement associated with this award for which reporting is required?

No

B.4 WHAT OPPORTUNITIES FOR TRAINING AND PROFESSIONAL DEVELOPMENT HAS THE PROJECT PROVIDED?

NOTHING TO REPORT

B.5 HOW HAVE THE RESULTS BEEN DISSEMINATED TO COMMUNITIES OF INTEREST?

NOTHING TO REPORT

B.6 WHAT DO YOU PLAN TO DO DURING THE NEXT REPORTING PERIOD TO ACCOMPLISH THE GOALS?

We plan to continue development to support the large NIH-funded imaging genetic initiatives. Specifically, the empirical genetic analyses developed this period will be expanded to large genomic studies across diverse cohorts. We will be concentrating on updating our website with videos on the use and how-to documentations on starting genetic imaging analyses. The accelerated version of the SOLAR will form the basis for the genetic-calculator web-site – now in development to support research by ENIGMA, HCP and other large collaborations.

We made significant progress along all four specific aims Aims. Activity on Aim 1 included in continues released development version of SOLAR-Eclipse (SE). In total, eleven incremental releases were made, each constituting a additional functionality, bug fixes and performance optimization change. All distribution was made through the NITRIC website https://www.nitrc.org/projects/se linux/. NITRIC registered 13242 downloads for our tool and lists it as the top 22rd most downloaded resource. The current production version of SE is used for performing imaging genetic research at several centers, including UCLA, University of South California, Yale University and Human Connectom Project at Washington University. Several novel and important developments were accomplished. Overall, the research performed during this year was described in 20 peer-reviewed publications including publications in JAMA psychiatry, Biological Psychiatry, Molecular Psychiatry, Neuroimage, Human Brain Mapping and others. To-date, over 150 manuscripts acknowledged support of this R01. Dr. Kochunov used the momentum generated by this application to secure NIH funding for a large hybrid CPU/GPU computational cluster funded through a shared instrumentation grant (S10 OD023696 01A1). This cluster will lay the foundation for high-performance computing for the next generation of genetic imaging grants.

Two chief progress directions were continued development described in Specific Aims 1 and 3: accelerating genetic analyses for high-throughput imaging genetics applications and development of empirical approaches for measuring relatedness. The acceleration of additive genetic variance calculations that were published by our group in the last two years [1-3] is now complemented by accelerated GWAS approach. The manuscript describing this approach was accepted for publication in Nature Communications (impact factor =12.1). It is "Fast and Powerful Genome Wide Association Analysis of Dense Genetic Data with High Dimensional Imaging Phenotypes" and describes an approximation-based inference testing that can accelerate the GWAS analysis in related individuals by a factor of 10³. This approach is now implemented in the SOLAR-Eclipse using both CPU and GPU implementations. Implementation of this algorithm on GPU provided an additional 1,000 fold improvement when used on a K-80 GPU computer at the Human Connectome Project/ Washington University. We are now in the process of writing a manuscript tentatively titled "Accelerating imaging genetics approaches using algorithms and graphics processing units" where we will formally evaluate the accuracy and performance of accelerated and classical methods. This project will be performed in collaboration with HCP using the newly released genotyping data for this consortium. The second focus was on development of empirical approaches for measuring relatedness directly from the high-density genetic data as described in Specific Aim 3. Quantitative genetic analyses rely on the estimates of relatedness or shared genetic variance among subjects: coefficients of relationship (CR). CR can be inferred from self-reported degrees of relatedness to other study participants or estimated empirically using genome-wide scans of single nucleotide polymorphisms (SNP). In the first study we hypothesized that the empirical CR constructed using whole genome genotyping information may provide more accurate measurements of shared genetic variance among study participants drawn from the same geographical area when compared to self-reported CR. We evaluated the performance of 12 state-of-the-art pairwise relatedness inference methods using a data set with 2485 individuals contained in several large pedigrees that span up to six generations. We observed that all methods have high accuracy (92-99%) when detecting first-, second-and third-degree relationships. Their accuracy dwindle for the more distant relationships. The identical by descent (IBD) segment-based methods however the long running time 24-80 hours reduced their practicality[4]. Based on this evaluation, we chose two methods that provided the good CR accuracy/performance benchmark: The Kinship-based INference for Genome wide association study (KING) method was developed to closely approximate the self-reported CR values. It is frequently used to verify the self-reported relationships in family samples [5]. A second approach, the Weighted Allelic Correlation (WAC) approach, was developed to study "missing

heritability" of complex phenotypes; this refers to the fact that heritability values for some traits may appear to be lower studies in studies of unrelated individuals rather than family or twinbased studies [6]. Both methods were integrated natively in SOLAR-Eclipse. They take raw genotype files in plink format and produce a pedigree file that can read by SOLAR-Eclipse using "load pedi" command. We partnered with HCP to test the hypothesis that that heritability estimates obtained from chromosomal CR values may provide additional information on the genetic contribution to the variance in complex traits [7, 8]. We tested these hypotheses in a large dataset of seventeen quantitative brain-related traits from four phenotypic domains collected by HCP. We observed that the whole-genome heritability estimates were significantly higher (p<0.001) using empirical relationships than these obtained based on self-reports. WAC with weighting on minor allele frequency produced the highest average heritability ($p < 10^{-6}$) estimates. Partitioning of shared variance into genetic and environmental components gave results that were independent of the CR approach. Chromosomal heritability estimates (the proportion of heritability arising from each chromosome) were significantly correlated with the length of the chromosome ($r \sim 0.7$). The patterns of per-chromosome heritability values were similar in traits from the same domain, among the traits that had significant genetic correlations and among the traits that described similar biological value but were not genetically correlated. Our findings suggest complex polygenic inheritance for quantitative traits. The manuscript describing the findings are now in submission to NeuroImage. The pedigree structures generated in these analyses are available through NITRC.

The work in Aim 2 included continued discussion with the International Neuroinformatics Coordinating Facility's Neuroimaging Data Sharing Task Force on the standardizing the imaging genetic format. At present our team is evaluating several of the proposed data format solution. We have tested the HDF5 based format in the parallel computing environment and found it to perform well for both reading and writing large volumes of data and parallel access from multiple servers. In addition, SOLAR-Eclipse team participated in the development of a standardized rsFMRI pipeline for genetic analyses. We published the preliminary heritability study describing good agreement among additive genetic variance obtained in two independent cohorts[9]. We also developed a web-based 3D viewing software to demonstrate results of large collaborative genetic studies[10].

Progress on Aim 4 has led to acknowledgement of SOLAR-Eclipse in publications in high impact journals, including PNAS, Nature, JAMA Psychiatry, Biological Psychiatry, Neuroimage, Human Brain Mapping and others[11-30].

References

- 1. Ganjgahi, H., et al., *Fast and powerful heritability inference for family-based neuroimaging studies.* Neuroimage, 2015.
- Winkler, A.M., et al., *Faster permutation inference in brain imaging*. Neuroimage, 2016. 141: p. 502-516.
- 3. Winkler, A.M., et al., *Multi-level block permutation*. Neuroimage, 2016. **123**: p. 253-268.
- 4. Ramstetter, M.D., et al., *Benchmarking Relatedness Inference Methods with Genome-Wide Data from Thousands of Relatives*. Genetics, 2017. **207**(1): p. 75-82.
- 5. Manichaikul, A., et al., *Robust relationship inference in genome-wide association studies*. Bioinformatics, 2010. **26**(22): p. 2867-2873.
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- 8. Visscher, Peter M., et al., *Genome Partitioning of Genetic Variation for Height from* 11,214 Sibling Pairs. American Journal of Human Genetics, 2007. **81**(5): p. 1104-1110.
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- 10. Zhang, G., et al., *ENIGMA-Viewer: interactive visualization strategies for conveying effect sizes in meta-analysis.* BMC Bioinformatics, 2017. **18**(Suppl 6): p. 253.
- 11. Kochunov, P., et al., *Diffusion-weighted imaging uncovers likely sources of processing-speed deficits in schizophrenia.* Proc Natl Acad Sci U S A, 2016. **113**(47): p. 13504-13509.
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 111: p. 300-311.
- 15. Kochunov, P., et al., *The common genetic influence over processing speed and white matter microstructure: Evidence from the Old Order Amish and Human Connectome Projects*. Neuroimage, 2015. **125**: p. 189-197.
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- 18. Sprooten, E., et al., A comprehensive tractography study of patients with bipolar disorder and their unaffected siblings. Human Brain Mapping, 2016. **37**(10): p. 3474-3485.
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- Rowland, L.M., et al., Medial Frontal GABA is Lower in Older Schizophrenia: A MEGA-PRESS with Macromolecule Suppression Study. Molecular psychiatry, 2016. 21(2): p. 198-204.
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- 22. Kuehner, R.M., et al., *Cognitive profiles and heritability estimates in the Old Order Amish.* Psychiatric Genetics, 2016. **26**(4): p. 178-183.
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24.	Unpublished
25.	Unpublished
26.	Kochunov, P., et al., Integration of routine OA data into mega†analysis may improve

Kochunov, P., et al., Integration of routine QA data into mega†analysis may improve quality and sensitivity of multisite diffusion tensor imaging studies. Human Brain Mapping, 2018. 39(2): p. 1015-1023.

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- Jahanshad, N., et al., Multi-site genetic analysis of diffusion images and voxelwise heritability analysis: A pilot project of the ENIGMA-DTI working group. Neuroimage, 2013. doi:pii: S1053-8119(13)00408-4. 10.1016/j.neuroimage.2013.04.061.

C.1 PUBLICATIONS

Are there publications or manuscripts accepted for publication in a journal or other publication (e.g., book, one-time publication, monograph) during the reporting period resulting directly from this award?

Yes

Publications Reported for this Reporting Period

Public Access Compliance	Citation
Complete	 Stein JL, Medland SE, Vasquez AA, Hibar DP, Senstad RE, Winkler AM, Toro R, Appel K, Bartecek R, Bergmann Ø, Bernard M, Brown AA, Cannon DM, Chakravarty MM, Christoforou A, Domin M, Grimm O, Hollinshead M, Holmes AJ, Homuth G, Hottenga JJ Langan C, Lopez LM, Hansell NK, Hwang KS, Kim S, Laje G, Lee PH, Liu X, Loth E, Lourdusamy A, Mattingsdal M, Mohnke S, Maniega SM, Nho K, Nugent AC, O'Brien C, Papmeyer M, Pütz B, Ramasamy A, Rasmussen J, Rijpkema M, Risacher SL, Roddey JC, Rose EJ, Ryten M, Shen L, Sprooten E, Strengman E, Teumer A, Trabzuni D, Turner J, van Eijk K, van Erp TG, van Tol MJ, Wittfeld K, Wolf C, Woudstra S, Aleman A, Alhusaini S, Almasy L, Binder EB, Brohawn DG, Cantor RM, Carless MA, Corvin A, Czisch M, Curran JE, Davies G, de Almeida MA, Delanty N, Depondt C, Duggirala R, Dyer TD, Erk S, Fagerness J, Fox PT, Freimer NB, Gill M, Göring HH, Hagler DJ, Hoehn D, Holsboer F, Hoogman M, Hosten N, Jahanshad N, Johnson MP, Kasperaviciute D, Kent JW Jr, Kochunov P, Lancaster JL, Lawrie SM, Liewald DC, Mandl R, Matarin M, Mattheisen M, Meisenzahl E, Melle I, Moses EK, Mühleisen TW, Nauck M, Nöthen MM, Olvera RL, Pandolfo M, Pike GB, Puls R, Reinvang I, Rentería ME, Rietschel M, Roffman JL, Royle NA, Rujescu D, Savitz J, Schnack HG, Schnell K, Seiferth N, Smith C, Steen VM, Valdés Hernández MC, Van den Heuvel M, van der Wee NJ, Van Haren NE, Veltman JA, Völzke H, Walker R, Westlye LT, Whelan CD, Agartz I, Boomsma DI, Cavalleri GL, Dale AM, Djurovic S, Drevets WC, Hagoort P, Hall J, Heinz A, Jack CR Jr, Foroud TM, Le Hellard S, Macciardi F, Montgomery GW, Poline JB, Porteous DJ, Sisodiya SM, Starr JM, Sussmann J, Toga AW, Veltman DJ, Walter H, Weiner MW, Bis JC, Ikram MA, Smith AV, Gudnason V, Tzourio C, Vernooij MW, Launer LJ, DeCarli C, Seshadri S, Andreassen OA, Apostolova LG, Bastin ME, Blangero J, Brunner HG, Buckner RL, Cichon S, Coppola G, de Zubicaray GI, Deary IJ, Donohoe G, de Geus EJ, Espeseth T, Fernández G, Glahn DC, Grabe HJ, Hardy J, Hulshoff Pol HE, Jenkinson M, Kahn RS, McDo
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N/A: Not Peer Reviewed	Unpublished
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Complete	 Whelan CD, Altmann A, Botía JA, Jahanshad N, Hibar DP, Absil J, Alhusaini S, Alvim MKM, Auvinen P, Bartolini E, Bergo FPG, Bernardes T, Blackmon K, Braga B, Caligiuri ME, Calvo A, Carr SJ, Chen J, Chen S, Cherubini A, David P, Domin M, Foley S, França W, Haaker G, Isaev D, Keller SS, Kotikalapudi R, Kowalczyk MA, Kuzniecky R, Langner S, Lenge M, Leyden KM, Liu M, Loi RQ, Martin P, Mascalchi M, Morita ME, Pariente JC Rodríguez-Cruces R, Rummel C, Saavalainen T, Semmelroch MK, Severino M, Thomas RH, Tondelli M, Tortora D, Vaudano AE, Vivash L, von Podewils F, Wagner J, Weber B, Yao Y, Yasuda CL, Zhang G, Bargalló N, Bender B, Bernasconi N, Bernasconi A, Bernhardt BC, Blümcke I, Carlson C, Cavalleri GL, Cendes F, Concha L, Delanty N, Depondt C, Devinsky O, Doherty CP, Focke NK, Gambardella A, Guerrini R, Hamandi K, Jackson GD, Kälviäinen R, Kochunov P, Kwan P, Labate A, McDonald CR, Meletti S, O'Brien TJ, Ourselin S, Richardson MP, Striano P, Thesen T, Wiest R, Zhang J, Vezzani A, Ryten M, Thompson PM, Sisodiya SM. Structural brain abnormalities in the common epilepsies assessed in a worldwide ENIGMA study. Brain : a journal of neurology. 2018 February 1;141(2):391-408. PubMed PMID: 29365066; PubMed Central PMCID: PMC5837616. 				
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PMC Journal - In process	 Kelly S, Jahanshad N, Zalesky A, Kochunov P, Agartz I, Alloza C, Andreassen OA, Arango C, Banaj N, Bouix S, Bousman CA, Brouwer RM, Bruggemann J, Bustillo J, Cahn W, Calhoun V, Cannon D, Carr V, Catts S, Chen J, Chen JX, Chen X, Chiapponi C, Cho KK, Ciullo V, Corvin AS, Crespo-Facorro B, Cropley V, De Rossi P, Diaz-Caneja CM, Dickie EW, Ehrlich S, Fan FM, Faskowitz J, Fatouros-Bergman H, Flyckt L, Ford JM, Fouche JP, Fukunaga M, Gill M, Glahn DC, Gollub R, Goudzwaard ED, Guo H, Gur RE, Gur RC, Gurholt TP, Hashimoto R, Hatton SN, Henskens FA, Hibar DP, Hickie IB, Hong LE, Horacek J, Howells FM, Hulshoff Pol HE, Hyde CL, Isaev D, Jablensky A, Jansen PR, Janssen J, Jönsson EG, Jung LA, Kahn RS, Kikinis Z, Liu K, Klauser P, Knöchel C, Kubicki M, Lagopoulos J, Langen C, Lawrie S, Lenroot RK, Lim KO, Lopez- Jaramillo C, Lyall A, Magnotta V, Mandl RCW, Mathalon DH, McCarley RW, McCarthy- Jones S, McDonald C, McEwen S, McIntosh A, Melicher T, Mesholam-Gately RI, Michie PT, Mowry B, Mueller BA, Newell DT, O'Donnell P, Oertel-Knöchel V, Oestreich L Paciga SA, Pantelis C, Pasternak O, Pearlson G, Pellicano GR, Pereira A, Pineda Zapata J, Piras F, Potkin SG, Preda A, Rasser PE, Roalf DR, Roiz R, Roos A, 				

Rotenberg D, Satterthwaite TD, Savadjiev P, Schall U, Scott RJ, Seal ML, Seidman LJ, Shannon Weickert C, Whelan CD, Shenton ME, Kwon JS, Spalletta G, Spaniel F, Sprooten E, Stäblein M, Stein DJ, Sundram S, Tan Y, Tan S, Tang S, Temmingh HS, Westlye LT, Tønnesen S, Tordesillas-Gutierrez D, Doan NT, Vaidya J, van Haren NEM, Vargas CD, Vecchio D, Velakoulis D, Voineskos A, Voyvodic JQ, Wang Z, Wan P, Wei D, Weickert TW, Whalley H, White T, Whitford TJ, Wojcik JD, Xiang H, Xie Z, Yamamori H, Yang F, Yao N, Zhang G, Zhao J, van Erp TGM, Turner J, Thompson PM, Donohoe G. Widespread white matter microstructural differences in schizophrenia across 4322 individuals: results from the ENIGMA Schizophrenia DTI Working Group. Molecular
psychiatry. 2018 May;23(5):1261-1269. PubMed PMID: 29038599.

C.2 WEBSITE(S) OR OTHER INTERNET SITE(S)

Category	Explanation			
Research Material, Research Material, Research Material	http://solar-eclipse-genetics.org/ is the main site for solar eclipse including manual page.			

C.3 TECHNOLOGIES OR TECHNIQUES

NOTHING TO REPORT

C.4 INVENTIONS, PATENT APPLICATIONS, AND/OR LICENSES

Have inventions, patent applications and/or licenses resulted from the award during the reporting period? No

If yes, has this information been previously provided to the PHS or to the official responsible for patent matters at the grantee organization? No

C.5 OTHER PRODUCTS AND RESOURCE SHARING

Category	Explanation				
Software	http://www.nitrc.org/projects/se_linux/ Is the main distribution point for the solar-eclipse software. It also provides the github link for source code and version references				
Research Material	http://solar-eclipse-genetics.org/ is the main site for solar eclipse including manual page.				
Data or Databases	http://enigma-viewer.org/ is the site for reporting results from large mega-analytical cohorts analyzed using solar-eclipse				

D. PARTICIPANTS

Commons ID	S/K	Name	Degree(s)	Role	Cal	Aca	Sum	Foreign	Country	SS
		Tunio	Dog.00(0)	TROID	ou.	7100	Cum	Org	country	
KOCHUNOV	Y	KOCHUNOV, PETER V	MS,MS,P HD	PD/PI	EFFORT					NA
Glossary of acro S/K - Senior/Key DOB - Date of B Cal - Person Mo Aca - Person Mo Sum - Person M	y Birth onths (Ca onths (A	cademic)				SS - Ši RE - Re DI - Div OT - Of	pplement S entry Supp ersity Supp	lement lement	tion Affiliation	1
D.2 PERSONNEL		TES								
D.2.a Level of Eff										
Will there be, in th or the PD/PI(s) o minimum amount	r other s	senior/key perso	nnel designa	ted in the No	5% or more in otice of Award	the level d, or (2) a	of effort from reduction in	n what was a the level of e	pproved by the fort below the	ne ager ne
No										
D.2.b New Senio	r/Key Pe	ersonnel								
D.2.b New Senio Are there, or will t			personnel?							
			personnel?							
Are there, or will t No	here be,	, new senior/key	personnel?							
Are there, or will t No D.2.c Changes in	here be	, new senior/key Support		f senior/key	personnel sir	ice the las	t reporting p	period?		
Are there, or will t No	here be	, new senior/key Support		f senior/key	personnel sir	ice the las	t reporting p	period?		
Are there, or will t No D.2.c Changes in Has there been a No	here be, Other S change	, new senior/key Support in the active oth		f senior/key	personnel sir	ice the las	t reporting p	period?		
Are there, or will t No D.2.c Changes in Has there been a No D.2.d New Other	here be, Other S change Significa	, new senior/key Support in the active oth ant Contributors	ner support o		personnel sir	ice the las	t reporting p	period?		
Are there, or will t No D.2.c Changes in Has there been a No D.2.d New Other Are there, or will t	here be, Other S change Significa	, new senior/key Support in the active oth ant Contributors	ner support o		personnel sir	ice the las	t reporting p	period?		
Are there, or will t No D.2.c Changes in Has there been a No D.2.d New Other Are there, or will t	here be, Other S change Significa	, new senior/key Support in the active oth ant Contributors , new other signi	ner support o		personnel sir	ice the las	t reporting p	period?		
Are there, or will t No D.2.c Changes in Has there been a No D.2.d New Other Are there, or will t	here be, Other S change Significa	, new senior/key Support in the active oth ant Contributors , new other signi	ner support o		personnel sir	ice the las	t reporting p	period?		
Are there, or will t No D.2.c Changes in Has there been a No D.2.d New Other Are there, or will t	here be, Other S change Significa here be, PI) Lead	, new senior/key Support in the active oth ant Contributors , new other signi	ificant contrib	outors?		ice the las	t reporting p	period?		

E. IMPACT

E.1 WHAT IS THE IMPACT ON THE DEVELOPMENT OF HUMAN RESOURCES?

Not Applicable

E.2 WHAT IS THE IMPACT ON PHYSICAL, INSTITUTIONAL, OR INFORMATION RESOURCES THAT FORM INFRASTRUCTURE?

NOTHING TO REPORT

E.3 WHAT IS THE IMPACT ON TECHNOLOGY TRANSFER?

Not Applicable

E.4 WHAT DOLLAR AMOUNT OF THE AWARD'S BUDGET IS BEING SPENT IN FOREIGN COUNTRY(IES)?

NOTHING TO REPORT

F. CHANGES

F.1 CHANGES IN APPROACH AND REASONS FOR CHANGE

Not Applicable

F.2 ACTUAL OR ANTICIPATED CHALLENGES OR DELAYS AND ACTIONS OR PLANS TO RESOLVE THEM

NOTHING TO REPORT

F.3 SIGNIFICANT CHANGES TO HUMAN SUBJECTS, VERTEBRATE ANIMALS, BIOHAZARDS, AND/OR SELECT AGENTS

F.3.a Human Subjects

No Change

F.3.b Vertebrate Animals

No Change

F.3.c Biohazards

No Change

F.3.d Select Agents

No Change

G. SPECIAL REPORTING REQUIREMENTS

G.1 SPECIAL NOTICE OF AWARD TERMS AND FUNDING OPPORTUNITIES ANNOUNCEMENT REPORTING REQUIREMENTS

NOTHING TO REPORT							
G.2 RESPONSIBLE CONDUCT OF RESEARCH							
Not Applicable							
G.3 M	G.3 MENTOR'S REPORT OR SPONSOR COMMENTS						
Not A	Not Applicable						
G.4 H	G.4 HUMAN SUBJECTS						
G.4.a	Does the project involve hu	man subjects?					
No							
	Inclusion Enrollment Data						
	oplicable						
G.4.c	ClinicalTrials.gov						
Does	this project include one or m	nore applicable clinic	cal trials that must be	registered in ClinicalTrials.gov under FDAAA?			
G.5 H	UMAN SUBJECTS EDUCA	TION REQUIREME	NT				
Are th	ere personnel on this projec	t who are newly inv	olved in the design o	r conduct of human subjects research?			
G.6 H	UMAN EMBRYONIC STEM	CELLS (HESCS)					
Does funde	Does this project involve human embryonic stem cells (only hESC lines listed as approved in the NIH Registry may be used in NIH funded research)?						
No							
G.7 V	G.7 VERTEBRATE ANIMALS						
Does	this project involve vertebra	te animals?					
No							
G.8 P	G.8 PROJECT/PERFORMANCE SITES						
	Organization Name:	DUNS	Congressional District	Address			
	Primary: University of Maryland, Baltimore	188435911	MD-07	UNIVERSITY OF MARYLAND BALTIMORE PO Box 21247 Baltimore MD 21228			
	University of Maryland, Baltimore	188435911	MD-007	UNIVERSITY OF MARYLAND BALTIMORE Office of Research and Development Baltimore MD 212011508			
	University of Texas Health Science Center at San Antonio800772162TX-0217703 Floyd Curl Drive, Mail Code 7828 San Antonio TX 782293900						

RPPR

University of Warwick	231745683	00-000	University House Kirby Corner Road Coventry, West Midlands
University of Maryland Baltimore	188435911	MD-007	Maryland Psychiatric Research Center Grounds of Spring Grove Hospital Center Baltimore MD 212284663
UNIVERSITY OF MARYLAND, BALTIMORE, OFFICE OF RESEARCH AND DEVELOPMENT	188435911		UNIVERSITY OF MARYLAND BALTIMORE 620 W LEXINGTON ST, 4TH FL BALTIMORE MD 212011508
University of Maryland, Baltimore	188435911	MD-07	UNIVERSITY OF MARYLAND BALTIMORE PO Box 21247 Baltimore MD 212011508
University of Maryland, Baltimore	188435911	MD-007	UNIVERSITY OF MARYLAND BALTIMORE Office of Research and Development Baltimore MD 212011508
University of Texas Health Science Center at San Antonio	800772162	TX-021	7703 Floyd Curl Drive, Mail Code 7828 San Antonio TX 782293900
University of Warwick	231745683	00-000	University House Kirby Corner Road Coventry, West Midlands
University of Maryland Baltimore	188435911	MD-007	Maryland Psychiatric Research Center Grounds of Spring Grove Hospital Center Baltimore MD 212284663

G.9 FOREIGN COMPONENT

No foreign component

G.10 ESTIMATED UNOBLIGATED BALANCE

G.10.a Is it anticipated that an estimated unobligated balance (including prior year carryover) will be greater than 25% of the current year's total approved budget?

No

G.11 PROGRAM INCOME

Is program income anticipated during the next budget period?

No

G.12 F&A COSTS

Is there a change in performance sites that will affect F&A costs?

No

QVR NIH Business System (NBS) Accounting Details

PRINT DOWNLOAD CLOSE PI: KOCHUNOV, PETER V FY: 2018 Total IMPACII Award Amt: \$400,001

Obligation Details for Project:

5R01EB015611-06

External Organization:

UNIVERSITY OF MARYLAND BALTIMORE

Accounting System Totals

Type:	Subaccount:domestic(P)	
Award Document Number:	<u>REB015611B</u>	Click hyperlink for accounting details for all projects with this document number

TIMING INFORMATION: QVR gathers disbursement data from NBS/nVision on a nightly basis, however, PMS data in NBS may lag as much as 2 weeks.

IC	CAN	CAN Budget Obligated Dt Last NBS Obligated NBS Disbursed						
		FY		Disburse. Dt	\$	\$	Balance	
EB 8	3015183	3 2018	2018-08-08		\$ 400,001.00	\$ 0.00	\$400,001.00	

Accounting System Transactions

Accounting System Transactions									
IC	CAN	000	NBS Doc Num	NBS Transact. (Obligation Amt	Disbursement			
				Dt		Amt			
EB	8015183	414E	380REB015611B*10001	2018-08-08	\$ 400,001.00	\$ 0.00			
				Grand Totals:	\$ 400,001.00	\$ 0.00			