# Harvard Medical School Curriculum Vitae

Date Prepared: 04/18/2023

Name: Olga Demler

Office Address: Division of Preventive Medicine

Brigham and Women's Hospital / Harvard Medical School

900 Commonwealth Ave, Boston MA, 02215

Work Phone: (617) 278-0861

**Work Email:** odemler@bwh.harvard.edu

Work FAX: (617) 731-3843

**Education** 

1992 BS Pure Mathematics Novosibirsk State

University,

Novosibirsk, Russia

1998 MS Engineering-Economic Systems Stanford University

and Operations Research Stanford, CA

1999 MA Statistics and Applied Probability University of California at

Santa Barbara, CA

Boston, MA

2012 PhD Biostatistics Boston University (BU),

(Professor Ralph D'Agostino,

Professor Michael Pencina)

**Postdoctoral Training** 

**Faculty Academic Appointments** 

2/13-8/14 Research Fellow Department of Medicine, Brigham and Women's (Professor Nancy Cook) Hospital/Harvard Medical

Professor Nancy Cook) Hospital/Harvard Medical School (BWH/HMS),

School (BWI I/I II)

Boston, MA

9/14-7/18 Instructor Medicine Harvard Medical School,

Boston, MA

7/18-12/21 Assistant Professor Medicine Harvard Medical School,

Boston, MA

1/22-Assistant Professor, Medicine Harvard Medical School, Boston, MA part time present 1/22-Tenured Senior Computer Science Swiss Federal Institute of Scientist Research, Technology (ETH), present Zurich, Switzerland part time

# Appointments at Hospitals/Affiliated Institutions

2014-2022 Associate Division of Preventive Medicine Brigham and Women's Biostatistician (DPM) Hospital, Boston, MA Division of Preventive Medicine Brigham and Women's 2022-Associate Hospital, present Biostatistician, part (DPM) Boston, MA time

# **Honors and Prizes**

1999	Graduate Student Scholarship Award for Academic Achievements	Department of Statistics and Applied Probability, the University of California at Santa Barbara, Santa Barbara, CA	Academic achievements
2008	Mu Sigma Rho, the National Statistics Honor Society	Boston Chapter of the American Statistical Association, Boston, MA	Academic achievements in graduate biostatistics program at Boston University
2017	Chair's Research Award	Department of Medicine Brigham and Women's Hospital	2017 Research Performance Incentive Program
2020	Alan M. Lerner young investigator research Award	Cardiovascular Division, Brigham and Women's Hospital	Won the award for our work on "Heterogeneous Effects of Omega-3 (n-3) Downstream Fatty Acids (FAs) and Oxylipins on Risk of Cardiovascular Disease: Results of Two Metabolomics Substudies of the Randomized Vitamin D and Omega-3 Trial (VITAL)."

# Report of Funded and Unfunded Projects

# **Funding Information**

**Past** 

2013-2016 Novel applications of risk reclassification methods in cardiovascular disease risk prediction

National Institutes of Health (NIH)/National Heart, Lung and Blood Institute (NHLBI) R01 HL113080

Statistician (PI: Cook)

The goal of this project is to develop new methods for comparing risk prediction models, particularly those that are related to reclassification calibration. It will extend these methods to other study designs and settings other than binary outcomes.

2013-2017

High density lipoprotein (HDL) heterogeneity and function, statin therapy, and cardiovascular disease (CVD) outcomes

NIH / NHLBI 1R01HL117861

Statistician (PI: Mora)

The major goal of this project is to evaluate HDL particle heterogeneity and function in relation to randomized statin treatment and prospectively ascertained CVD outcomes in the Justification for the Use of Statin in Prevention: An Intervention Trial Evaluating Rosuvastatin (JUPITER) and Treating to New Targets (TNT) clinical trials.

2014-2015

HDL heterogeneity and function, statin therapy, and CVD outcomes (supplement) NIH / NHLBI 1R01HL117861 (supplement 1)

Statistician (PI: Mora)

The goal of this project is to use state-of-the-art lipidomics as a tool to more specifically and comprehensively identify lipid biomarkers of normal and dysfunctional HDL.

2016-2020

Inflammatory mediators, cardiovascular health, and longevity in women NIH/NHLBI R01 AG051654

Statistician (PI: Cheng)

The major goal of this study is to advance our understanding of how upstream inflammatory pathways are related to healthy cardiovascular aging and longevity in women. To achieve this goal, metabolomic profiling will be performed on baseline blood samples of 5,000 participants from the Women's Health Study (WHS) to study associations between cardiovascular and general morbidity outcomes and eicosanoid mediators of systemic inflammation.

2020-2021

Predicting the Severity and Cardiovascular Complications of COVID-19 and the Risk of ACE Inhibitors/ARBs using Machine Learning

Co-PI - (\$100,000)

The goal of this project is to identify phenotypes for the severity of COVID-19 clinical presentation, with a particular emphasis on cardiovascular manifestations, using electronic health data.

2018-2022

PROMINENT (The Pemafibrate to Reduce cardiovascular OutcoMes by reducing triglycerides IN diabetic patiENTs)

Associate Statistician at Data Coordinating Center (PI: Ridker)

The Triglyceride Reduction and Acute Cardiovascular Events (TRACE) trial is a randomized, double blind, placebo-controlled, international trial evaluating the ability of the potent PPAR-alpha agonist, K-877, to reduce rates of myocardial infarction, stroke, unstable angina requiring unplanned revascularization, and cardiovascular death in among 10,000 men and women with type 2 diabetes (T2D).

2017-2022

Coronary Heart Disease (CHD) Risk and Metabolomic Profiles of Discordant Lipids

PI – (\$809,386 - total direct costs for entire project period beginning in 2017)

This project is designed to combine statistical learning/data mining methods with results of my prior methodological research on impact of correlation on risk prediction models in order to find the most informative combinations of an extended panel of lipid parameters that best

predict CHD events and determine metabolomic profiles of individuals with various lipid combinations.

2019-2022 Flexible Framework for Cardiovascular Risk Prediction in a Clinical Setting American Heart Association Methods Validation Grant 17IGMV33860009

PI – (\$184,868 – direct costs)

This project is designed to create a Flexible Framework of cardiovascular risk prediction models by combining existing validated risk prediction models such as Pooled Cohort equations, QRISK3 and MESA with rich longitudinal person-level data available in Partners EHR system.

#### Current

2021-2023 Machine learning risk stratification in patients with ASCVD: A personalized approach NIH/NHLBI R21 HL156174

Co-I (PI: Mora) – (\$268,514)

The goal of this study is to apply new artificial intelligence methods to develop a free and publicly available computer program that will calculate each patient's long-term chance of having or dying from a repeat heart attack or stroke based on his or her individual risk profile.

2019-2023 Total Plasma and IgG Glycomes, Statin Therapy and ASCVD events

NIH/NHLBI R01 HL117861

Co-I (PI:Mora)-(\$3,404,424)

The goal of this study is to advance our understanding of the human glycome by identifying glycosylation patterns related positively or inversely to incident ASCVD and risk factors, in particular inflammatory and vascular risk factors.

2021-2024 Plasma Metabolite and Proteome Signatures for Migraine Classification

NIH R61 NS122074

Co-I (Pls: Chasman, Rist) (\$2,065,041)

This proposal seeks to identify and validate plasma-based metabolite and protein signatures for migraine with the goal of improving current approaches to migraine treatment and identifying new therapeutic targets.

2022-2026 Targeting the Active Resolution of Inflammation for Cardiovascular Disease Prevention

1 R01 HL160799

Co-I (PI: Mora, S) (\$3,404,424)

The aim of this proposal is to comprehensively examine determinants of circulating levels of specialized pro-resolving mediators (SPMs) and their relationships to the resolution or promotion of inflammation and with future CVD risk.

2022-2026 Rheumatoid Arthritis and the Risk of Cardiovascular Disease: Biomarkers, Risk Prediction, and Underlying Mechanisms

R01 HL163580

Co-I (PI: Solomon, D).(\$3,138,853)

The overarching goal of this proposal is to identify protein biomarkers for cardiovascular disease (CVD) in patients with rheumatoid arthritis (RA), leveraging the structure of a randomized controlled trial and rigorous methods for deriving and validating a risk score.

2022-2027 Decoding mechanisms underlying metabolic dysregulation in obesity and digestive cancer risk

U01 CA272452

Co-I (PI: Tobias D. / Giovannuci, E. / Zhang, X) (\$7,287,767)

The project aims to identify new protein biomarkers and protein signatures for type 2 diabetes, using modern proteomics techniques to comprehensively assess a large set of the proteins circulating in the bloodstream in order to better understand the complex relationships between proteins, metabolites, diet, lifestyle, and type 2 diabetes.

2023-2024 Consensus Framework for Cardiovascular Risk Prediction in a Clinical Setting R21HL167173

PI: Demler (\$268,500)

The goal of this project is to evaluate ways to combine published risk prediction models in real-life clinical setting using Machine Learning models such as Super Learner and eXtreme Gradient Boosting in clinical settings and develop a novel method called the Consensus Framework. This novel method has the consensus property because it combines multiple published and validated risk models to ensure not only good overall performance but also good performance in important subgroups of patients. The Consensus Framework is adapted to clinical practice because it can handle limited information or additional risk factors. We will also assess specific properties of prognostic risk prediction and how they inform the selection of the most appropriate class of models.

2023-2025 Integrating medical image data and assessments for personalized cardiovascular risk estimation (Dataspectrum4CVD)

Swiss Data Science Center/PHRT

PI: Demler (CHF286.000)

This project involves integration of full spectrum of data modalities (genetic, raw image data and biomarkers) with novel Machine Learning and Deep Learning approaches in order to improve performance of prognostic models for ASCVD and diagnostic models for coronary artery calcification (CAC) score.

## **Pending**

2024-2025 Using machine learning/artificial intelligence methods to improve ASCVD risk estimates for misclassified low-risk patients using image, genetic and -omics data

R21 NIH NHLBI

PI: Demler (\$492,250)

The goal of this project is to improve the risk classification of low-risk individuals who constitute up to 30% of those who later develop CVD by integrating the full spectrum of data in UKBiobank and MGB EHR data: we will integrate genetic, image and biomarker data. To achieve this we will develop a diagnostic deep learning model of the CAC score from chest X-rays and other images and additionally, we will to replicate existing published highly accurate machine learning models that use ECG images to extract rich phenotypic data that is not available from traditional human reads.

# Report of Local Teaching and Training

#### **Teaching of Students in Courses**

**Harvard University Courses** 

Advanced Statistical Learning/Data Harvard T.H. Chan School of Public 2016

Mining Methods Health 4-hour course

Dr. Donna Spiegelman's research group: Graduate Students, Postdocs, Research

Associates

2018 Metabolomic Studies: from Design Stage Harvard T.H. Chan School of Public

to the Validation Stage Graduate Students enrolled in HDSC 325 course

Health. Guest Lecturer Health Data Science Capstone Course HDSC 325

2022	A Crash Course on Convolutional Neural Networks	Harvard T.H. Chan School of Public Health graduate course. Guest Lecturer Introduction to Machine Learning and Risk Prediction EPI288

**Other Teaching** 

2010 Introduction to Statistical Programming Boston University, School of Public

Using SAS, lecturer Health,
Boston, MA

Graduate students 4-hr lectures and 1-hr practice sessions

per wk for 12 wks

2012 Introduction to Biostatistics I Clinical and Translational Science

graduate program, Tufts University,

Boston, MA

Masters and PhD students (most with MD

degree)

5-hr lectures and office hours per wk for

20 wks

# Formal Teaching of Peers (e.g., CME and other continuing education courses)

No presentations below were sponsored by outside entities.

2014-2017	Introduction to Predictive Modeling Center for Clinical Investigation (CCI), Brigham and Women's Hospital	One 4-hour lecture yearly Boston, MA
2018	A Crash Course in Statistical Learning Methods Center for Clinical Investigation (CCI), Brigham and Women's Hospital	One 4-hour lecture Boston, MA
2022	A Three-Day Crash Course in Machine Learning Methods, Center for Clinical Investigation, Brigham and Women's Hospital	Three 3-hour lectures Boston, MA

## **Local Invited Presentations**

No presentations below were sponsored by outside entities.

2012	Improvement of Area Under the Receiver Operating Characteristics Curve (AUC), Significance of New Predictors and Limits of Validity of Some Common AUC Tests Used in Risk Prediction Modeling/ Research Design Center/Biostatistics Research Center Seminar Tufts University Medical School
2012	Impact of New Variables on Discrimination in Risk Prediction Models/invited seminar series lecture Division of Preventive Medicine, BWH
2013	Impact of Correlation on Predictive Ability of a Biomarker/Division Seminar Division of Preventive Medicine, BWH

2014	Improvement of AUC for Risk Prediction Models/colloquium Statistical Methods in Epidemiology Working Group Harvard TH Chan School of Public Health
2015	Tests of Calibration and Goodness-of-Fit in the Survival Setting/colloquium at Statistical Methods in Epidemiology Working Group Harvard TH Chan School of Public Health
2018	Asymptotic distribution of $\Delta AUC$ , NRIs, and IDI based on theory of U-statistics /colloquium at Statistical Methods in Epidemiology Working Group Harvard TH Chan School of Public Health
2020	Heterogeneous Effects of Omega-3 Downstream Fatty Acids and Oxylipins on Risk of Cardiovascular Disease: Results of Two Metabolomics Substudies of the Randomized Vitamin D and Omega-3 Trial (VITAL) /Alan M. Lerner Research Symposium, Division of Cardiovascular Medicine, Brigham and Women's Hospital
2020	Building Research Cohort from Partner's Electronic Health Records data /Division of Preventive Medicine Work-in-Progress Seminar
2022	Focus on Disease Mechanisms: Biostatistical Perspective / Division of Preventive Medicine Work-in-Progress Seminar

# Report of Regional, National and International Invited Teaching and Presentations

No presentations below were sponsored by outside entities.

# **National**

2011	Misuse of DeLong Test to Compare AUCs of Nested Models/ Oral Presentation Miami Beach, Florida (Joint Statistical Meetings 2011)
2011	Comparing AUCs of Nested Models/ Oral Presentation World cardiovascular, diabetes, and obesity online conference (Target Meeting 1st World Cardiovascular, Diabetes, and Obesity Online Conference 2011)
2012	Equivalence of Improvement in Area Under ROC Curve and Linear Discriminant Analysis Coefficient Under Assumption of Normality/ Oral Presentation San Diego, California (Joint Statistical Meetings 2012)
2013	Impact of Correlation on Predictive Ability of a Biomarker/ Oral Presentation Montreal, Canada (Joint Statistical Meetings 2013)
2015	Methods for reclassification calibration in the survival setting/ Oral Presentation Seattle, Washington (Joint Statistical Meetings 2015)
2016	Asymptotic Distribution of ΔAUC, NRI, and IDI Based on U-Statistics Theory/ Oral Presentation Chicago, Illinois (Joint Statistical Meetings 2016)

2017	Asymptotic Distribution of ΔAUC, NRI, and IDI Based on U-Statistics Theory/ Oral Presentation
	Washington, DC (Eastern North American Regional (ENAR) Meeting of International Biometrics Society 2017)
2017	Measures of Predictive Model Performance and Event Rate: How Much They Vary and How to Make Them Comparable across Studies / Oral Presentation Baltimore, MD (Joint Statistical Meetings 2017)
2018	Powering Biomarker Discovery Studies for Training and Validation / Oral Presentation Vancouver, Canada (Joint Statistical Meetings 2018)
2019	AUC as a Measure of the Probability of Benefit in the Context of Randomized Controlled Trials / Oral Presentation Denver, Colorado (Joint Statistical Meetings 2019)
2019	Effects of Marine Omega-3 Supplementation on Fatty Acids and Bioactive Lipids and Associations with Risk of Cardiovascular Disease: Secondary Analysis of the Vital Trial Philadelphia, Pennsylvania (AHA Scientific Sessions 2019)
2019	Anti-inflammatory HDL Function and Incident Cardiovascular/Death Events: A Secondary Analysis of the JUPITER Trial Boston, Massachusetts (BWH Women in Science Symposium)
2020	WiNN: Drift Correction by White Noise Guided Normalization for Metabolomic Studies that does not Rely on Quality Control Samples Online (16th International Conference of the Metabolomics Society)
2020	Effects of Omega-3 (n-3) Supplementation on Downstream Fatty Acids (FAs) and Oxylipins and Risk of Cardiovascular Disease: Results of Two Metabolomics Substudies of the Randomized VITamin D and OmegA-3 TriaL Online (American Heart Association Scientific Meetings)
2020	WiNN: Drift Correction by White Noise Normalization for Metabolomic Studies Online (Joint Statistical Meetings)
2021	Using Stochastic Orders to Evaluate Performance of Predictive Models: Intransitivity, Area under the ROC and Strength of Stochastic Order Relationships Online Topic-Contributed Invited Presentation (Joint Statistical Meetings)
2022	Asymptotic properties of AUC under the null in the training-test setting: assessing AUC change for polygenic risk scores and machine learning risk prediction models. (Joint Statistical Meetings)
2022	Glycan biomarkers for cardiovascular events (3rd Meeting of the Human Glycome Project, An International Conference of Glycomics Society. Split, Croatia)
2023	Non-transitivity of Area Under the Receiver Operating Characteristics Curve (Conference of Health Inference and Learning, Boston MA)
2023	AUC estimator for nested matched case-control studies (Joint Statistical Meetings, Toronto Canada)

# Report of Technological and Other Scientific Innovations

Implemented novel goodness-of-fit test in survival setting in SAS and R (2015)

SAS and R code are available online at http://ncook.bwh.harvard.edu/r-code.html and in the Appendix of the manuscript which has been published in *Statistics in Medicine* journal (Research Investigation #19)

SARS2: COVID risk score of

hospitalization and

death

SARS2 model is provided as a web interface for seamless calculation of the risk scores and risk categories. https://dashti.bwh.harvard.edu/sars2/

# **Report of Scholarship**

#### Peer-reviewed publications in print or other media

#### Research investigations

- 1. Wang PS, **Demler OV**, Kessler RC. The adequacy of treatment for serious mental illness in the United States. American Journal of Public Health. Am J Public Health. 2002;92(1):92-98. PMID: 11772769. Cited by 435.
- Kessler RC, Andrade LH, Bijl RV, Offord DR, **Demler OV**, Stein DJ. The effects of co-morbidity on the onset and persistence of generalized anxiety disorder in the ICPE surveys. International Consortium in Psychiatric Epidemiology. *Psychol Med*. 2002;32(7):1213-1225. PubMed PMID: 12420891. Cited by 79.
- Kessler RC, Ormel J, Demler OV, Stang PE. Comorbid mental disorders account for the role impairment of commonly occurring chronic physical disorders: results from the National Comorbidity Survey. *J Occup Environ Med*. 2003;45(12):1257-1266. PubMed PMID: 14665811. Cited by 291.
- 4. Kessler RC, Berglund P, **Demler OV**, Jin R, Koretz D, Merikangas KR, Rush AJ, Walters EE, Wang PS. National Comorbidity Survey Replication. The epidemiology of major depressive disorder: results from the National Comorbidity Survey Replication (NCS-R). *JAMA*. 2003 18;289(23):3095-30105. PubMed PMID: 12813115. Cited by 6780.
- Kessler RC, Berglund P, Chiu WT, **Demler OV**, Heeringa S, Hiripi E, Jin R, Pennell BE, Walters EE, Zaslavsky A, Zheng H. The US National Comorbidity Survey Replication (NCS-R): design and field procedures. *Int J Methods Psychiatr Res.* 2004;13(2):69-92. PubMed PMID: 15297905. Cited by 643.
- 6. Kessler RC, Abelson J, Demler OV, Escobar JI, Gibbon M, Guyer ME, Howes MJ, Jin R, Vega WA, Walters EE, Wang P, Zaslavsky A, Zheng H. Clinical calibration of DSM-IV diagnoses in the World Mental Health (WMH) version of the World Health Organization (WHO) Composite International Diagnostic Interview (WMHCIDI). *Int J Methods Psychiatr Res.* 2004;13(2):122-39. PubMed PMID: 15297907. Cited by 380.
- 7. Kessler RC, **Demler OV**, Frank RG, Olfson M, Pincus HA, Walters EE, Wang P, Wells KB, Zaslavsky AM. Prevalence and treatment of mental disorders, 1990 to 2003. *N Engl J Med*. 2005

- Jun 16;352(24):2515-23. PubMed PMID: 15958807; PubMed Central PMCID: PMC2847367. Cited by 1497.
- Kessler RC, Chiu WT, Demler OV, Merikangas KR, Walters EE. Prevalence, severity, and comorbidity of 12-month DSM-IV disorders in the National Comorbidity Survey Replication. *Arch Gen Psychiatry*. 2005;62(6):617-627. PubMed PMID: 15939839; PubMed Central PMCID: PMC2847357. Cited by 9228.
- 9. Kessler RC, Berglund P, **Demler OV**, Jin R, Merikangas KR, Walters EE. Lifetime prevalence and age-of-onset distributions of DSM-IV disorders in the National Comorbidity Survey Replication. *Arch Gen Psychiatry*. 2005;62(6):593-602. PubMed PMID: 15939837. Cited by 11988
- 10. Kessler RC, Birnbaum H, **Demler OV**, Falloon IR, Gagnon E, Guyer M, Howes MJ, Kendler KS, Shi L, Walters E, Wu EQ. The prevalence and correlates of nonaffective psychosis in the National Comorbidity Survey Replication (NCS-R). *Biol Psychiatry*. 2005;58(8):668-676. Epub 2005 Jul 14. PubMed PMID: 16023620; PubMed Central PMCID: PMC2847859. Cited by 233.
- 11. Kessler RC, Adler L, Ames M, **Demler OV**, Faraone S, Hiripi E, Howes MJ, Jin R, Secnik K, Spencer T, Ustun TB, Walters EE. The World Health Organization Adult ADHD Self-Report Scale (ASRS): a short screening scale for use in the general population. *Psychol Med.* 2005;35(2):245-56. PubMed PMID: 15841682. Cited by 1178.
- 12. Wang PS, **Demler OV**, Olfson M, Pincus HA, Wells KB, Kessler RC. Changing profiles of service sectors used for mental health care in the United States. *Am J Psychiatry*. 2006;163(7):1187-98. PubMed PMID: 16816223; PubMed Central PMCID: PMC1941780. Cited by 280.
- 13. Kessler RC, Adler L, Barkley R, Biederman J, Conners CK, **Demler OV**, Faraone SV, Greenhill LL, Howes MJ, Secnik K, Spencer T, Ustun TB, Walters EE, Zaslavsky AM. The prevalence and correlates of adult ADHD in the United States: results from the National Comorbidity Survey Replication. *Am J Psychiatry*. 2006;163(4):716-23. PubMed PMID: 16585449; PubMed Central PMCID: PMC2859678. Cited by 2765.
- Demler OV, Pencina MJ, D'Agostino RB Sr. Equivalence of improvement in area under ROC curve and linear discriminant analysis coefficient under assumption of normality. Stat Med. 2011;30(12):1410-8. doi: 10.1002/sim.4196. Epub 2011 Feb 21. PubMed PMID: 21337594. Cited by 28.
- 15. **Demler OV**, Pencina MJ, D'Agostino RB Sr. Misuse of DeLong test to compare AUCs for nested models. *Stat Med.* 2012;31(23):2577-87. doi: 10.1002/sim.5328. Epub 2012 Mar 13. PubMed PMID: 22415937; PubMed Central PMCID: PMC3684152. Cited by 81.
- 16. Pencina MJ, D'Agostino RB Sr, **Demler OV**. Novel metrics for evaluating improvement in discrimination: net reclassification and integrated discrimination improvement for normal variables and nested models. *Stat Med.* 2012;31(2):101-13. doi: 10.1002/sim.4348. Epub 2011 Dec 7. PubMed PMID: 22147389; PubMed Central PMCID: PMC3341978. Cited by 121.
- 17. **Demler OV**, Pencina MJ, D'Agostino RB Sr. Impact of correlation on predictive ability of biomarkers. *Stat Med*. 2013;32(24):4196-4210. doi: 10.1002/sim.5824. PubMed PMID: 23640729. Cited by 6.
- 18. White KT, Moorthy MV, Akinkuolie AO, **Demler OV**, Ridker PM, Cook NR, and Mora S. Identifying an optimal cutpoint for the diagnosis of hypertriglyceridemia in the nonfasting state. *Clinical chemistry* 2015;61(9):1156-63. doi: 10.1373/clinchem.2015.241752. Cited by 11.

- 19. **Demler OV**, Paynter NP, Cook NR Tests of Calibration and Goodness-of-Fit in the Survival Setting. *Stat Med* 2015 34(10), 1659-1680. doi: 10.1002/sim.6428. Cited by 22.
- 20. **Demler OV**, Pencina MP, Cook NR, D'Agostino RB Sr. Asymptotic Distribution of ΔAUC, NRIs, and IDI based on Theory of U-statistics. *Stat Med*. 2017 36(21);334-60.
- 21. Khera AV\*, **Demler OV**\*, Adelman SJ, Collins HL, Glynn RJ, Ridker PM, Rader DJ, Mora S Cholesterol Efflux Capacity, HDL Particle Number, and Incident Cardiovascular Events. *Circulation* 2017;135(25):2494-2504. doi: 10.1161/CIRCULATIONAHA.116.025678. PubMed PMID: 28450350; PubMed Central PMCID: PMC5490983. (\*Drs. Khera and Demler contributed equally to this work). Cited by 3.
- 22. Harada PHN, **Demler OV**, Dugani SB, Akinkuolie AO, Moorthy MV, Ridker PM, Cook NR, Pradhan AD, Mora S. Lipoprotein insulin resistance score and risk of incident diabetes during extended follow-up of 20 years: The Women's Health Study. *J Clin Lipidol*. 2017;11(5), 1257-1267.e2. doi: 10.1016/j.jacl.2017.06.008. PubMed PMID: 28733174; PubMed Central PMCID: PMC5644504.
- 23. **Demler OV**, Paynter NP, Cook NR. Reclassification calibration test for censored survival data: performance and comparison to goodness-of-fit criteria. Diagnostic and prognostic research. 2018 Dec;2(1):16. doi: 10.1186/s41512-018-0034-5.
- 24. Tobias DK, Lawler PR, Harada PH, **Demler OV**, Ridker PM, Manson JE, Cheng S, Mora S. Circulating Branched-Chain Amino Acids and Incident Cardiovascular Disease in a Prospective Cohort of US Women. Circulation. Genomic and precision medicine. 2018 April;11(4):e002157. PubMed PMID: 29572205; PubMed Central PMCID: PMC5880282.
- 25. Farukhi ZM, **Demler OV**, Caulfield MP, Kulkarni K, Wohlgemuth J, Cobble M, Luttmann-Gibson H, Li C, Nelson JR, Cook NR, Buring JE. Comparison of Nonfasting and Fasting Lipoprotein Subfractions and Size in 15,397 Apparently Healthy Individuals: An Analysis from the VITamin D and OmegA-3 TriaL (VITAL). Journal of Clinical Lipidology. 2020 Feb 21.
- 26. Luttmann-Gibson H, Mora S, Camargo CA, Cook NR, **Demler OV**, Ghoshal A, Wohlgemuth J, Kulkarni K, Larsen J, Prentice J, Cobble M, Bubes V, Li C, Friedenberg G, Lee IM, Buring JE, Manson JE. Serum 25-hydroxyvitamin D in the VITamin D and OmegA-3 TriaL (VITAL): Clinical and demographic characteristics associated with baseline and change with randomized vitamin D treatment. Contemp Clin Trials. 2019 Dec;87:105854. doi: 10.1016/j.cct.2019.105854. Epub 2019 Oct 24. PubMed PMID: 31669447; PubMed Central PMCID: PMC6875603.
- Enserro DM, Demler OV, Pencina MJ, D'Agostino RB Sr. Measures for evaluation of prognostic improvement under multivariate normality for nested and nonnested models. Stat Med. 2019 Sep 10;38(20):3817-3831. doi: 10.1002/sim.8204. Epub 2019 Jun 18. PubMed PMID: 31211443; PubMed Central PMCID: PMC6827341.
- 28. Antonelli J, Claggett BL, Henglin M, Kim A, Ovsak G, Kim N, Deng K, Rao K, Tyagi O, Watrous JD, Lagerborg KA, Hushcha PV, **Demler OV**, Mora S, Niiranen TJ, Pereira AC, Jain M, Cheng S. Statistical Workflow for Feature Selection in Human Metabolomics Data. Metabolites. 2019 Jul 12;9(7). doi: 10.3390/metabo9070143. Review. PubMed PMID: 31336989; PubMed Central PMCID: PMC6680705.
- 29. Henglin M, Niiranen T, Watrous JD, Lagerborg KA, Antonelli J, Claggett BL, Demosthenes EJ, von Jeinsen B, **Demler O**, Vasan RS, Larson MG, Jain M, Cheng S. A Single Visualization Technique for Displaying Multiple Metabolite-Phenotype Associations. Metabolites. 2019 Jul

- 2;9(7). doi: 10.3390/metabo9070128. PubMed PMID: 31269707; PubMed Central PMCID: PMC6680673.
- 30. Ahmad S, Moorthy MV, **Demler OV**, Hu FB, Ridker PM, Chasman DI, Mora S. Assessment of Risk Factors and Biomarkers Associated With Risk of Cardiovascular Disease Among Women Consuming a Mediterranean Diet. JAMA Netw Open. 2018 Dec 7;1(8):e185708. doi: 10.1001/jamanetworkopen.2018.5708. PubMed PMID: 30646282; PubMed Central PMCID: PMC632432.
- 31. Kessler RC, Bauer MS, Bishop TM, **Demler OV**, Dobscha SK, Gildea SM, Goulet JL, Karras E, Kreyenbuhl J, Landes SJ, Liu H. Using Administrative Data to Predict Suicide After Psychiatric Hospitalization in the Veterans Health Administration System. Frontiers in Psychiatry. 2020 May 6;11:390.
- 32. Chasman DI, Giulianini F, **Demler OV**, Udler MS. Pleiotropy-Based Decomposition of Genetic Risk Scores: Association and Interaction Analysis for Type 2 Diabetes and CAD. The American Journal of Human Genetics. 2020 Apr 16.
- Dashti H, Westler WM, Wedell JR, Demler OV, Eghbalnia HR, Markley JL, Mora S. Probabilistic identification of saccharide moieties in biomolecules and their protein complexes. Sci Data. 2020 Jul 3;7(1):210. doi: 10.1038/s41597-020-0547-y. PubMed PMID: 32620933; PubMed Central PMCID: PMC7335193.
- 34. Sels D, Dashti H, Mora S, **Demler OV**, Demler E. Quantum approximate Bayesian computation for NMR model inference. Nat Mach Intell. 2020 Jul;2(7):396-402. doi: 10.1038/s42256-020-0198-x. Epub 2020 Jul 6. PubMed PMID: 33163858; PubMed Central PMCID: PMC7643990.
- 35. Ajala ON\*, **Demler OV**\*, Liu Y, Farukhi Z, Adelman SJ, Collins HL, Ridker PM, Rader DJ, Glynn RJ, Mora S. Anti-Inflammatory HDL Function, Incident Cardiovascular Events, and Mortality: A Secondary Analysis of the JUPITER Randomized Clinical Trial. J Am Heart Assoc. 2020 Sep;9(17):e016507. doi: 10.1161/JAHA.119.016507. Epub 2020 Aug 15. PubMed PMID: 32799709; PubMed Central PMCID: PMC7660788.
- 36. **Demler OV**, Liu, Y, Luttmann-Gibson, H, Watrous, JD, Lagerborg, KA, Dashti, H, ... & Buring, JE., Manson, JE, Cheng, S, Jain M, Mora, S. (2020). One-Year Effects of Omega-3 Treatment on Fatty Acids, Oxylipins, and Related Bioactive Lipids and Their Associations with Clinical Lipid and Inflammatory Biomarkers: Findings from a Substudy of the Vitamin D and Omega-3 Trial (VITAL). Metabolites, 10(11), 431. PMID: 33120862
- 37. Dashti H, Roche EC, Bates DW, Mora S, **Demler OV**. SARS2 simplified scores to estimate risk of hospitalization and death among patients with COVID-19. Sci Rep. 2021 Mar 2;11(1):4945. doi: 10.1038/s41598-021-84603-0. PMID: 33654180.
- Dugani SB, Moorthy MV, Li C, **Demler OV**, Alsheikh-Ali AA, Ridker PM, Glynn RJ, Mora S. Association of Lipid, Inflammatory, and Metabolic Biomarkers With Age at Onset for Incident Coronary Heart Disease in Women. JAMA Cardiol. 2021 Apr 1;6(4):437-447. doi: 10.1001/jamacardio.2020.7073. PubMed PMID: 33471027; PubMed Central PMCID: PMC7818181.
- 39. Kessler RC, Bauer MS, Bishop TM, Bossarte RM, Castro VM, **Demler OV**, Gildea SM, Goulet JL, King AJ, Kennedy CJ, Landes SJ, Liu H, Luedtke A, Mair P, Marx BP, Nock MK, Petukhova MV, Pigeon WR, Sampson NA, Smoller JW, Miller A, Haas G, Benware J, Bradley J, Owen RR, House S, Urosevic S, Weinstock LM. Evaluation of a Model to Target High-risk Psychiatric Inpatients for an Intensive Postdischarge Suicide Prevention Intervention. JAMA Psychiatry. 2023 Mar 1;80(3):230-240. doi: 10.1001/jamapsychiatry.2022.4634. PMID: 36652267

- 40. Birukov A, Plavša B, Eichelmann F, Kuxhaus O, Hoshi RA, Rudman N, Štambuk T, Trbojević-Akmačić I, Schiborn C, Morze J, Mihelčić M, Cindrić A, Liu Y, **Demler OV**, Perola M, Mora S, Schulze MB, Lauc G, Wittenbecher C. Immunoglobulin G N-Glycosylation Signatures in Incident Type 2 Diabetes and Cardiovascular Disease. Diabetes Care. 2022 Nov 1;45(11):2729-2736. doi: 10.2337/dc22-0833. PMID: 36174116
- 41. Hoshi RA, Liu Y, Luttmann-Gibson H, Tiwari S, Giulianini F, Andres AM, Watrous JD, Cook NR, Costenbader KH, Okereke OI, Ridker PM, Manson JE, Lee IM, Vinayagamoorthy M, Cheng S, Copeland T, Jain M, Chasman\* DI, **Demler\* OV**, Mora\* S. Association of Physical Activity With Bioactive Lipids and Cardiovascular Events. Circ Res. 2022 Aug 5;131(4):e84-e99. doi: 10.1161/CIRCRESAHA.122.320952. Epub 2022 Jul 19. PMID: 35862024
- 42. Henglin M, Claggett BL, Antonelli J, Alotaibi M, Magalang GA, Watrous JD, Lagerborg KA, Ovsak G, Musso G, **Demler OV**, Vasan RS, Larson MG, Jain M, Cheng S. Quantitative Comparison of Statistical Methods for Analyzing Human Metabolomics Data. Metabolites. 2022 Jun 4;12(6):519. doi: 10.3390/metabo12060519. PMID: 35736452

## Non-peer reviewed scientific or medical publications/materials in print or other media

# Reviews, chapters, monographs and editorials

- 1. Kessler RC, Chiu WT, Colpe L, **Demler OV**, Merikangas KR, Walters EE, Wang PS. The prevalence and correlates of serious mental illness (SMI) in the National Comorbidity Survey Replication (NCS-R). In Manderscheid RW, JT Berry JT (Eds.). Mental Health, United States, 2004. Rockville, MD: Substance Abuse and Mental Health Services Administration; 2006. pp. 134-148.
- 2. Kessler RC, Berglund PA, Chiu W-T, **Demler OV**, Glantz M, Lane MC, Jin R, Merikangas KR, Nock M, Olfson M, Pincus,HA, Walters EE, Wang PS, Wells KB. The National Comorbidity Survey Replication (NCS-R): Cornerstone in improving mental health and mental health care in the United States. In Kessler RC, Üstün TB (Eds.). The WHO World Mental Health Surveys: Global Perspectives on the Epidemiology of Mental Disorders. New York: Cambridge University Press; 2008. pp. 165-209.
- 3. Dashti HT, Bates D, Fiskio JM, Roche EC, Mora S, **Demler OV**. Clinical Characteristics and Severity of COVID-19 Disease in Patients from Boston Area Hospitals. medRxiv. 2020 Aug 4;. doi: 10.1101/2020.07.27.20163071. PubMed PMID: 32793930; PubMed Central PMCID: PMC7418757.

#### Letters to the Editor

 Cook NR, Demler OV, Paynter NP. Clinical Risk Reclassification at 10 Years. Stat Med. 2017; Dec 10;36(28):4498-502.

#### **Thesis**

**Demler OV**. Impact of New Variables on Discrimination of Risk Prediction Models [dissertation]. Boston (MA): Boston University; 2012.

# **Narrative Report**

I split time between Computer Science department at ETH Zurich and Division of Preventive Medicine at Brigham and Women's Hospital / Harvard Medical. At both am teaching graduate students and medical professionals.

Teaching Harvard learners is an important part of my everyday work. Over the years I have been teaching a three-part crash course in Machine Learning Methods at the Center for Clinical Investigation (CCI), Brigham and Women's Hospital. I am also mentoring Harvard medical and graduate students and fellows. I mentor two mentees at any given time. These weekly meetings cover how to conduct statistical analyses and the theoretical basis behind each method, selecting appropriate methods. address study design issues, and how to implement the planned analysis using statistical software tools. Each mentee works on an individual project based on a novel research idea using datasets stored at the Division. Mentees submit their results to scientific conferences which are successfully selected for presentations. Each mentee has produced at least one paper in peer-reviewed journal as a first author. Manuscript preparation is another opportunity for them to learn how to report scientific findings, use precise unambiguous language, and conduct reproducible scientific research. I have mentored Drs. Zareen Farukhi (MPH student at Harvard Chan School of Public Health), Remi Ajala (postdoctoral T32 fellow at Harvard Medical School/BWH), Hesam Dashti (postdoctoral T32 fellow at Harvard Medical School/BWH). Currently I am mentoring Aniket Zinzuwadia (third year Harvard Medical School student) and Drs. Rosangela Hoshi (postdoctoral fellow at Harvard Medical School/BWH) and Chirag Vyas (Instructor in Medicine at Harvard Medical School).

Additionally, I teach Harvard fellows and faculty the theory and practice of cutting edge statistical analysis used in recent publications at the Division of Preventive Medicine Journal Club. I have covered the following topics: meta-analysis, Mendelian Randomization analysis, analysis of metabolomics data, boosting methods (HGBoost), blockchain for medical research, and others. I typically give a lecture explaining how a given novel method works and demonstrate how it has been used in practice with a publication that we then discuss. I answer their questions and give comments. Currently Journal Club is attended by Harvard faculty from the Division of Preventive Medicine and Harvard postdoctoral fellows (Drs. Rikuta Hamaya, Rosangela Hoshi) and Instructor in Medicine Dr. Zareen Farukhi.

I am a guest lecturer for the Harvard Chan graduate course EPI288 "Introduction to Machine Learning and Risk Prediction" (Spring semester), where I am teaching Neural Networks.

My research interests are two-fold: 1) I do methodological and clinical research in risk prediction and 2) I oversee statistical analyses in biomarker discovery clinical studies including metabolomics and other high throughput -omics studies.

My methodological work in risk prediction is focused on properties of the commonly used measure of discrimination called AUC (or c-statistic) and on the development of novel goodness of fit tests in survival settings. My work in risk prediction has been recognized through five first-authored peer-reviewed publications, two of which were among the top ten most cited papers in the Statistics in Medicine journal in 2012-2013. As part of my clinical research in risk prediction, I am leading a project funded by the American Heart Association on building flexible predictive models using a large-scale database of Partners Electronic Medical Records data from several major Boston-area hospitals.

My research in biomarker discovery focuses on all stages of conventional, NMR, LCMS and LCMS2 metabolomic biomarker discovery studies, mostly in the cardiovascular field with the goal of facilitating the discovery of novel drug targets and interventions. My research includes the development of rigorous analytic workflows of the analysis of metabolomics data starting with quality control procedures, signal correction, and further statistical analysis of metabolomic studies. I was awarded a five-year career development award (K01) from the National Heart, Lung, and Blood Institute to investigate coronary heart disease risk and metabolic profiles of individuals with discordant lipids.

Additionally, I provide statistical leadership in the analysis of several randomized clinical trials and observational studies including the Framingham Heart Study, Women's Health Study, JUPITER trial, and the VITAminD and OmegA-3 Trial (VITAL). I also participate in the biannual DSMB meetings for PROMINENT - one of the ongoing multi-center clinical trials and provide statistical expertise for the DSMB reports.