



International Phase II/NAT workshop

2024

September 25-26

The Ohio State University 1645 Neil Ave, Columbus, Ohio

September 25-26, 2024

Hamilton Hall, Ohio State University Medical Center, 1645 Neil Ave, Columbus, Ohio 43210

All roundtable and platform sessions include onsite and virtual (Zoom) presentations and discussions.

Workshop registration is free of charge to all participants. Refreshments and meals are provided at no cost to the speakers and to a limited number of onsite participants. Anyone interested in presenting or attending (either onsite or virtually) should send their contact information to David W. Hein at david.hein@louisville.edu as soon as possible since the number of additional onsite attendees for whom meals will be provided at no cost may be limited.

Special thanks to the Society of Toxicology for funding, the University of Louisville for logistic support and to Ohio State University for logistic support and hosting the workshop.

Organizing Committee

- David W. Hein, University of Louisville, Louisville, Kentucky USA
- Sotiria Boukouvala, Democritus University of Thrace, Alexandroupolis, Greece
- Giannoulis Fakis, Democritus University of Thrace, Alexandroupolis, Greece

The workshop will immediately follow the Pharmacogenomics Global Research Network meeting scheduled for September 23-25 at Ohio State University. Meeting site: https://www.pgrn.org/event-5628011.

<u>Accommodation Recommendations:</u>

The Blackwell Hotel: 2110 Tuttle Park Place, to Hamilton Hall 0.9 miles - 19 minute walk Hyatt House: 633 W 5th Avenue, to Hamilton Hall 0.9 miles - 21 minute walk Aloft: 1295 Olentangy River Road, to Hamilton Hall 1.2 miles - 26 minute walk Springhill Suites: 1421 Olentangy River Road, to Hamilton Hall 1.0 miles - 23 minute walk The Graduate (recently acquired by Hilton, so Hilton rewards will apply here): 750 N High Street, to Hamilton Hall 1.7 miles 38 minute walk









September 25

Registration and refreshments (2:30-3:00 pm)

Welcome by **David W. Hein**; Introduction of Organizing Committee; funding acknowledgements; expressions of thanks; schedule updates; conference dinners; participant introductions

Conference Roundtable (3:00-5:30 pm)

- NAT nomenclature committee and PharmVar expert panel updates on NAT2 allele nomenclature
 - Sotiria Boukouvala and Georgia Papanikolaou, Democritus University of Thrace, Alexandroupolis, Greece
- CPIC update on NAT2 allele function
 - David W. Hein, University of Louisville School of Medicine, Louisville, Kentucky USA
- CPIC therapeutic recommendations for hydralazine therapy related to NAT2 phenotype
 - Michael T. Eadon, Indiana University School of Medicine, Indianapolis, Indiana USA
- The use of language related to human diversity in pharmacogenetics: The case of NAT2
 - Kyle B. Brothers, University of Louisville School of Medicine, Louisville, Kentucky USA
- Discussion regarding NAT nomenclature updates and recommendations
 - All workshop participants

Platform Session I: Diversity in *NAT2* and other pharmacogenes (6:00-6:30 pm)

- Evolution of human diversity in NAT2 and other pharmacogenes
 - o Estella S. Poloni, University of Geneva, Genève, Switzerland

Private Conference Dinner (7 – 10 pm; <u>The Columbus Fish Market</u>, 1245 Olentangy River Road, Columbus Ohio 43212)

September 26

Breakfast (8:00- 8:30 am)

- The polymorphic enzymes NAT2, GSTM1, and GSTT1 and urinary bladder cancer risk after the structural change in the local chemical industry
 - o Klaus Golka, Leibniz Research Centre for Working Environment and Human Factors at TU Dortmund (IfADo). Dortmund, Germany (via Zoom)

Platform Session II: NATs in chronic diseases (8:30 to 10:30 am)

- Phenotypic Distribution of Acetylation and Hydroxylation Profiles Based on NAT2, CYP2E1, CYP3A4, and CYP3A5 Genes Across Brazil: Implications for Adverse Drug Reactions in Leprosy
 - Adalberto R. Santos, Oswaldo Cruz Institute, Rio de Janeiro, Brazil (via Zoom)
- The role of arylamine N-acetyltransferases in chronic degenerative diseases: exploring their possible function in the immune system
 - o Diana Portales-Perez, Autonomous University of San Luis Potosí, Mexico
- Non-coding and intergenic genetic variants of human arylamine N-acetyltransferase 2 (NAT2) gene are associated with differential plasma lipid and cholesterol levels and cardiometabolic disorders

Kyung U. Hong, Western New England University, Springfield, Massachusetts, USA (via Zoom)

- Differential expression of NAT1 in hormone receptor positive vs. negative female breast cancer
 - o Giannoulis Fakis, Democritus University of Thrace, Alexandroupolis, Greece

Refreshment Break (10:30 am to 11:00 am)

Platform Session III: NAT2 allele definition and function (11:00 am to 12:15 pm)

- Single nucleotide and copy number variation at the NAT2 locus, assessed according to PharmVar criteria and nomenclature
 - Georgia Papanikolaou, Democritus University of Thrace, Alexandroupolis, Greece
- Differences in β-naphthylamine metabolism and toxicity in Chinese hamster ovary cell lines transfected with human CYP1A2 and NAT2*4, NAT2*5B or NAT2*7B Nacetyltransferase 2 haplotypes
 - o Mariam R. Habil, University of Louisville, Louisville, Kentucky, USA
- The role of N-acetyltransferase 1 in epithelial to mesenchymal transition of U87 cells
 - Luke A. Schroeder, University of Louisville School of Medicine, Louisville, Kentucky USA

Conference lunch and roundtable discussion of *NAT2* **allele function (12:15 – 1:30 pm)**

- The effect of the rs1799931 c.857G>A (p.Gly286Glu) polymorphism on Nacetyltransferase 2-mediated carcinogen metabolism and genotoxicity differs with heterocyclic amine exposure
 - o David W. Hein, University of Louisville, Louisville, Kentucky USA

Platform Session IV: NATs and mitochondrial function (1:30 – 2:30 pm)

- Stable isotope tracing reveals an altered fate of glucose in N-acetyltransferase 1 knockout breast cancer cells
 - James TF Wise, Louisiana State University, Baton Rouge, Louisiana, USA
- The arylamine N-acetyltransferases as therapeutic targets in metabolic diseases associated with mitochondrial dysfunction
 - o Rodney F. Minchin, University of Queensland, Brisbane, Australia (via Zoom)

Platform Session V: NAT functions in xenobiotic metabolism and beyond (2:30-5:30 pm)

- N-acetyltransferase 2 genetic polymorphism modifies genotoxic and oxidative damage from new psychoactive substances
 - Raul Salazar-Gonzalez, Discovery Life Sciences, Malden, Massachusetts, USA
- Assaying NATs of human and other primates
 - o Ioanna Stavrakaki, Democritus University of Thrace, Alexandroupolis, Greece (currently at University of Crete, Heraklion, Greece; via Zoom)
- Bacterial NATs in xenobiotic metabolism
 - Maria-Giusy Papavergi, Democritus University of Thrace, Alexandroupolis, Greece (currently at Harvard Medical School, Cambridge, Massachusetts, USA)
- Bacterial NATs in secondary metabolism
 - Dionysios Patriarcheas, Democritus University of Thrace, Alexandroupolis, Greece (currently at University of West Virginia, Morgantown West Virginia, USA)
- NATs in plant-pathogenic fungi
 - Anthony E. Glenn, US Department of Agriculture, Agricultural Research Service, Athens, Georgia, USA (via Zoom)
- Unraveling the evolutionary origins of microbial NATs: from transglutaminases to acetyltransferases, and from xenobiotic biotransformation to the biosynthesis of secondary metabolites
 - Sotiria Boukouvala, Democritus University of Thrace, Alexandroupolis, Greece
- A summary of antibody reagents against NATs all available to a good home!
 - o Edith Sim, University of Oxford, Oxford, UK (via Zoom)

Platform Session VI: UDP- UDP-glucuronosyltransferases (5:30 – 6:30 pm)

- Drug-drug interactions between cannabinoids and UGT-mediated metabolism of opioids and benzodiazepines
 - Philip Lazarus, State University of New York at Buffalo, Buffalo, New York, USA
- Role of N-glycosylation in the activity, function, and cellular localization of human uridine diphosphate glucuronosyltransferase 1A6 (UGT1A6)
 - Yuejian Liu, University of British Columbia, Vancouver, British Columbia, Canada (via Zoom)
- Comprehensive characterization of rat and mouse UDP-glucuronosyltransferases
 Yuji Ishii, Kyushu University, Fukuoka, Japan (via Zoom)

Private Conference Celebration and Dinner (7 – 10 pm; upstairs loft, Wine on High 789 North High Street, Short North Arts District, Columbus, Ohio 43215; dinner provided by Hubbard Grill).

Celebrating over 25 years of International N-acetyltransferase Workshops hosted in eight different countries around the world

First International Arylamine N-acetyltransferase Workshop October 22-24,1998; Kuranda, Australia

Second International Arylamine N-acetyltransferase Workshop October 5-6, 2001; Oxford, UK

Third International Arylamine N-acetyltransferase Workshop August 27-28, 2004; Vancouver, Canada

Fourth International Arylamine N-acetyltransferase Workshop September 14-16, 2007; Alexandroupolis, Greece

Fifth International Arylamine N-acetyltransferase Workshop September 1-3, 2010; Paris, France (Meeting website)

Sixth International Arylamine N-acetyltransferase Workshop October 4-6, 2013; Toronto, Canada (Meeting website)

<u>Seventh International Workshop on N-acetyltransferases</u> June 18-20, 2016; Trier, Germany (<u>Meeting website</u>)

<u>Eighth International Workshop on N-acetyltransferases</u> September 25-26, 2024; Columbus, Ohio USA*

*Rescheduled from April 1-3, 2020 in Louisville, Kentucky USA due to worldwide COVID19 pandemic.

Onsite Participants

Ammar Almarzoog

Kuwait Cancer Control Center Kuwait, Cityu, Kuwait

Kyle B. Brothers

University of Louisville School of Medicine Louisville, Kentucky USA

Sotiria Boukouvala

Democritus University of Thrace Alexandroupolis, Greece

Jerome Dixon

Virginia Commonwealth University Richmond, Virginia USA

Michael T. Eadon

Indiana University School Medicine Indianapolis, Indiana USA

Giannoulis Fakis

Democritus University of Thrace Alexandroupolis, Greece

Yelena Guttman

University of California, San Francisco San Francisco, California USA

Mariam R. Habil

University of Louisville School of Medicine Louisville, Kentucky USA

David W. Hein

University of Louisville School of Medicine Louisville, Kentucky USA

Philip Lazarus

State University of New York at Buffalo Buffalo, New York, USA

Georgia Papanikolaou

Democritus University of Thrace Alexandroupolis, Greece

Maria-Giusy Papavergi

Harvard University Cambridge, Massachusetts USA

Dionysios Patriarcheas

West Virginia University Morgantown, West Virginia USA

Estella S. Poloni

University of Geneva Geneve, Switzerland

Diana Portales-Perez

Autonomous University of San Luis Potosi San Luis Potosi, Mexico

Raul Salazar-Gonzalez

Discovery Life Sciences Malden, Massachusetts USA

Luke A. Schroeder

University of Louisville School of Medicine Louisville, Kentucky USA

Marcus Stepp

AmplifyBio Columbus, Ohio USA

Chonlaphat Sukasem

Mahidol University Salaya, Thailand

Bailey Tibben

St. Jude Children's Research Hospital Memphis, Tennessee USA

Michelle Whirl-Carrillo

Stanford University Stanford, California USA

James TF Wise

Louisiana State University Baton Rouge, Louisiana USA

Virtual Speakers

Adalberto R. Santos

Oswaldo Cruz Institute Rio de Janeiro, Brazil

Klaus Golka

Leibniz Research Centre for Working Environment and Human Factors at TU Dortmund (IfADo), Dortmund, Germany

Rodney F. Minchin

University of Queensland Brisbane, Australia

Ioanna Stavrakaki

University of Crete Heraklion, Greece

Anthony E. Glenn

US Department of Agriculture Agricultural Research Service Athens, Georgia, USA

Kyung U. Hong

Western New England University Springfield, Massachusetts USA

Edith Sim

University of Oxford Oxford, UK

Yuejian Liu

University of British Columbia Vancouver, Canada

Yuji Ishii

Kyushu University Fukuoka, Japan

Virtual Participants

Christina Aquilante

University of Colorado Denver, Colorado, USA

Dimitra Basdani

Democritus University of Thrace Alexandroupolis, Greece

Joshua Hood

University of Louisville School of Medicine Louisville, Kentucky, USA

Nicola Laurieri

University of Oxford Oxford, UK

Yeeming Lee

University of Colorado Denver, Colorado, USA

Marcia Quinhones Pires Lopes

Fiocruz Rio de Janeiro, Brazil

James Martin

University of Colorado Denver, Colorado, USA

Charlene McQueen

University of Arizona Tucson Arizona USA

Mederic Mouterde

University of Geneva Geneve, Switzerland

Andrew Paterson

University of Toronto Toronto. Canada

Mariana Puzer Regis

Fiocruz Rio de Janeiro, Brazil

Katrin Sangkuhl

Stanford University
Stanford, California USA

Victória de Moraes Silva

Fiocruz Rio de Janeiro, Brazil

Youssef Tawik

University of Mainz Mainz, Germany

Raquel Lima de Figueiredo Teixeira

Fiocruz Rio de Janeiro, Brazil

Janet Zang

US Food and Drug Administration College Park, Maryland USA

Sokratis Zekkas

Democritus University of Thrace Alexandroupolis, Greece

Phase II/N-acetyltransferase Workshop Photos

September 25-26, 2024 Columbus, Ohio, USA









