About OMICS Group

OMICS Group International is an amalgamation of Open Access publications and worldwide international science conferences and events. Established in the year 2007 with the sole aim of making the information on Sciences and technology 'Open Access', OMICS Group publishes 500 online open access scholarly journals in all aspects of Science, Engineering, Management and Technology journals. OMICS Group has been instrumental in taking the knowledge on Science & technology to the doorsteps of ordinary men and women. Research Scholars, Students, Libraries, Educational Institutions, Research centers and the industry are main stakeholders that benefitted greatly from this knowledge dissemination. OMICS International also organizes 500 International conferences annually across the globe, where knowledge transfer takes place through debates, round table discussions, poster presentations, workshops, symposia exhibitions.

About OMICS International Conferences

OMICS International is a pioneer and leading science event organizer, which publishes around 500 open access journals and conducts over 300 Medical, Clinical, Engineering, Life Sciences, Pharma scientific conferences all over the globe annually with the support of more than 1000 scientific associations and 30,000 editorial board members and 3.5 million followers to its credit.

OMICS International has organized 500 conferences, workshops and national symposiums across the major cities including San Francisco, Las Vegas, San Antonio, Omaha, Orlando, Raleigh, Santa Clara, Chicago, Philadelphia, Baltimore, United Kingdom, Valencia, Dubai, Beijing, Hyderabad, Bengaluru and Mumbai.

Keynote Address

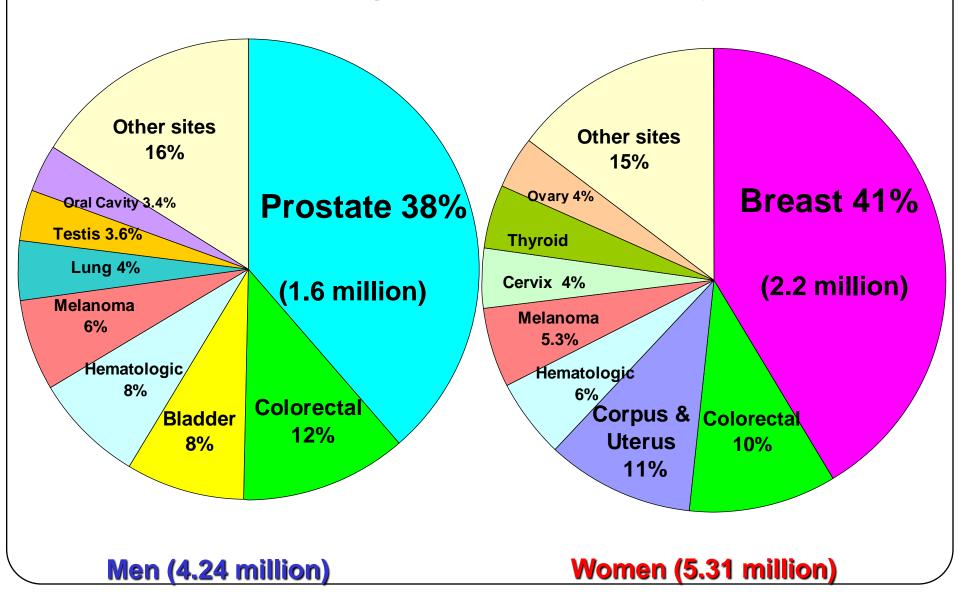
Metabolomics Research at NIH

Mukesh Verma

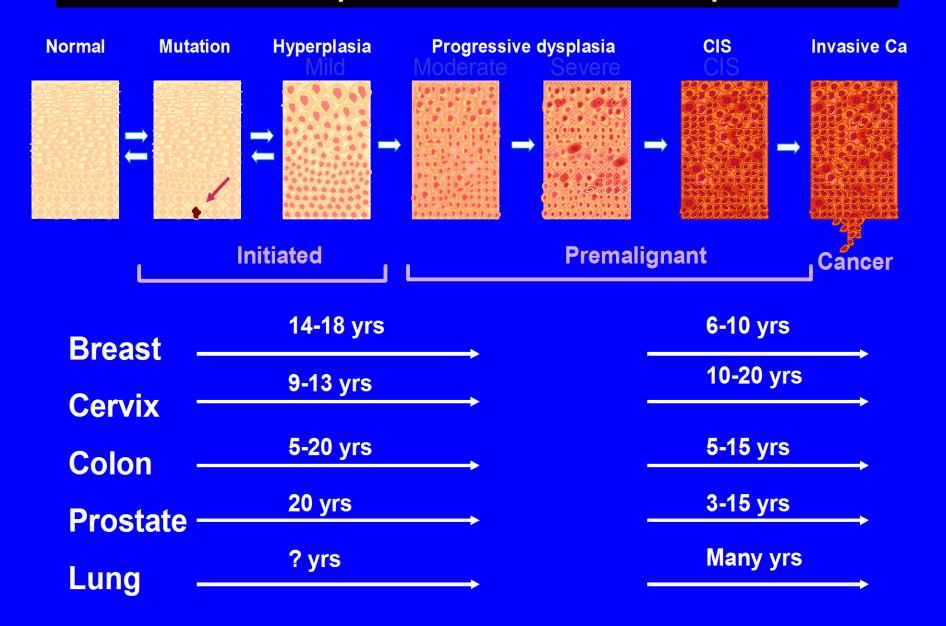
Chief, Methods and Technologies Branch
Program Director, Epidemiology and Genetics Research Program
Division of Cancer Control and Population Sciences
National Cancer Institute
National Institutes of Health

U.S. DEPARTMENT
OF HEALTH AND
N SERVICES
al Institutes
of Health

Estimated Number of Persons Alive in the U.S. Diagnosed with Cancer by Site



Cancer Development is a Multi-step Process



The Broad Reach of the NIH



NIH is an institution (Intramural Research)

- ~ 6,000 scientists
- ~ 10% of NIH budget

There are resources--

FY11 budget: \$31B

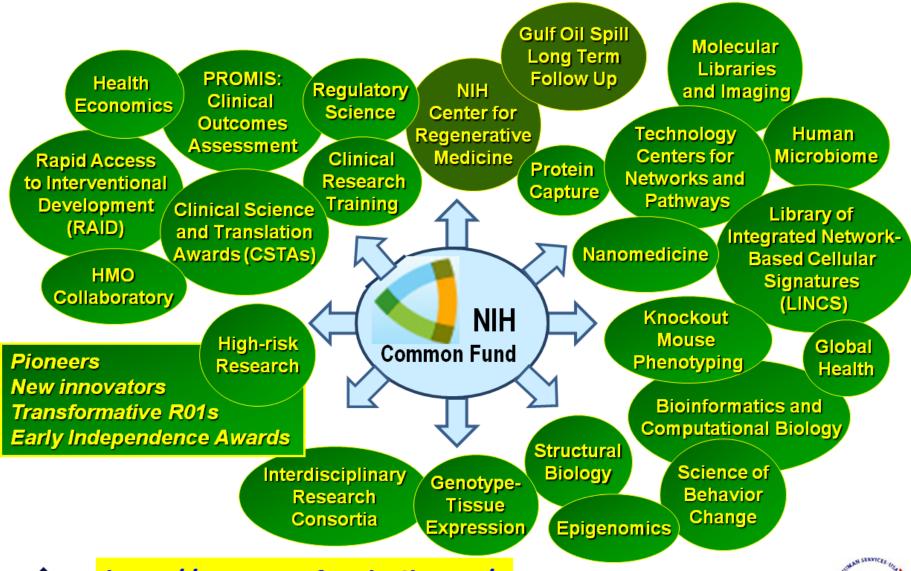


NIH *supports* institutions & people

(Extramural Research)

- > 4,000 institutions
- > 300,000 scientists & research personnel
- ~85% of the NIH budget

Cross-Cutting, trans-NIH Programs





http://commonfund.nih.gov/



NIH Common Funds: Metabolomics

Anderson, James (NIH/OD)
Smith, Philip (NIH/NIDDK)
Singer, Dinah (NIH/NCI)
Varmus, Harold (NIH/NCI)
Castle, Arthur (NIH/NIDDK)
Basavappa, Ravi (NIH/OD)
Wilder, Elizabeth (NIH/OD)
Kawazoe, Robin (NIH/OD)
Verma, Mukesh (NIH/NCI)

Potential Focus Area of the New Program (Varmus and Anderson)

- 1. Standards and protocols for measuring and quantifying most of the human metabolome.
- 2. Informatics tools to process, store and disseminate metabolomics information.
- A core understanding of how much and how often individual metabolites vary under normal conditions, especially in clinical samples.
- 4. Improved tools for measuring sub-cellular localization and changes in metabolite concentrations.

Metabolomics Working Group Members

Balshaw, David (NIEHS)	Castle, Arthur (NIDDK)	
Dutta, Chhanda (NIA)	Nadler, Laurie (NIMH)	
Okita, Richard (NIGMS)	Oversby, Steven (NEI)	
Proctor, Lita (NHGRI)	Shaughnessy, Daniel (NIEHS)	
Shum, Lillian (NIDCR)	Smith, Philip (NIDDK)	
Spalholz, Barbara (NCI)	Srinivas, Pothur R (NHLBI)	
Tagle, Danilo (NINDS)	Tigno, Xenia (NINR)	
Velazquez, Jose (NIA)	Verma, Mukesh (NCI)	
Winer, Karen (NICHD)	Zakhari, Samir (NIAAA)	

What is Metabolomics?

- Metabolomics enables <u>simultaneous identification</u> and <u>analysis</u> of multiple <u>metabolites</u> in cells, tissues and body fluids.
- Metabolomics deals with a <u>global analysis of small molecule</u> metabolites and <u>metabolic patterns</u>.
- Metabonomics, another term used in relation to <u>metabolic</u> <u>profiling</u>, refers specifically to <u>metabolic measurements</u> in organisms <u>in response to external stimuli</u> or <u>genetic modifications</u>.

(In the post genomic era, metabolomics has emerged as a field of great <u>significance</u> for both <u>translational</u> and <u>basic</u> biological <u>research</u>)

Additional Information about Metabolomics

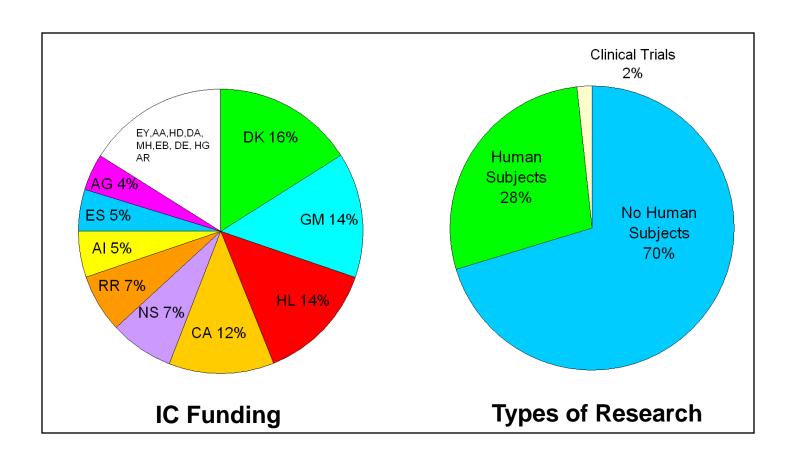
How individuals respond to therapy?

Metabolomics technologies permit <u>integration of</u> <u>biological pathways</u> to understand how living organisms <u>interact with its environment</u>

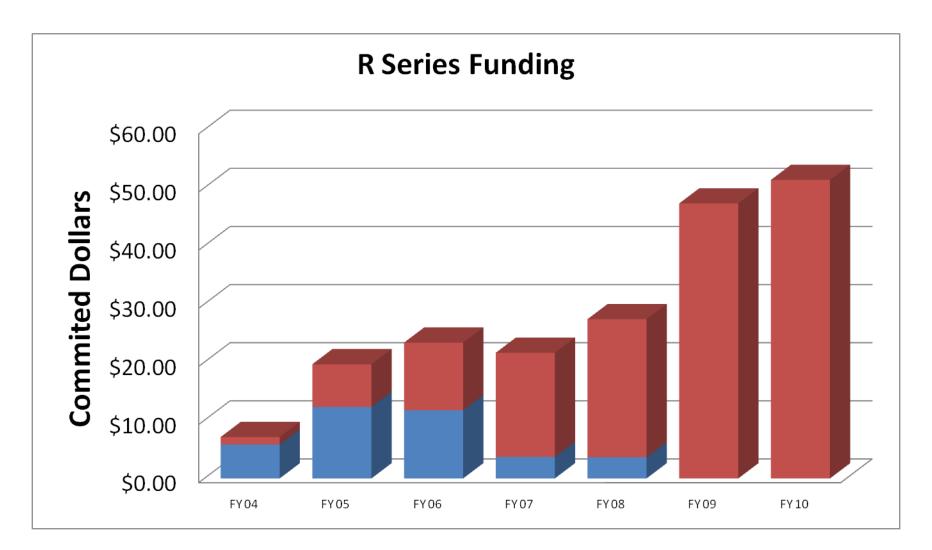
and

produce a <u>metabolic fingerprint</u> or 'metabolome', analogous to the genome or the proteome.

Distribution of Awarded Grants in Metabolomics (RPGs)



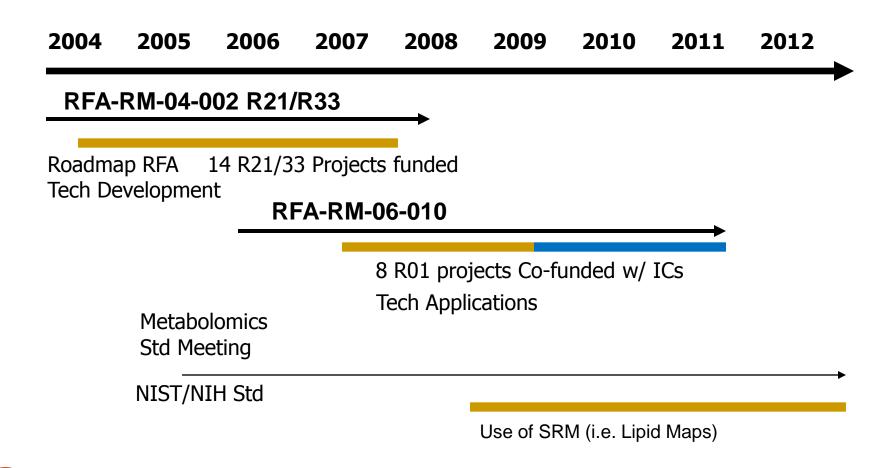
Growth of NIH Metabolomics Funding



Currently Funded Projects

- In 2010, at least 115 R series grants have self identified themselves with metabolomics in at least one specific aim.
- In 2010, 55 P or U series grants and cooperative agreements have at least one core or subproject self identified as metabolomics
- In most of these grants/agreements <u>metabolomics is a minor</u> <u>subcomponent</u> of overall project
- Most are <u>not technology development</u> or clinical/translational research (mostly discovery)

Metabolomics Roadmap Timeline



Awarded Projects (R21/R33)

Wayne R Matson	"Integrating LCEC/LCMS in a Single Metabolomics Platform"
Herbert Hill	"The Potential of Ion Mobility Mass Spectrometry for Metabolic Profiling"
Julian Griffin	"Metabolomics and Metabolic Compartmentation in the Brain"
Truman R Brown	"Metabolic Patterns in 1H NMR Spectra of Biofluids"
Jonathan V Sweedler	"Technologies for Cellular Neurometabolomics"
Michael R Sussman	"Isotope-Assisted Differential Metabolomics"
Fred E Regnier	"Tools for Comparative Metabolomics"
Gerhard Wagner	"An Integrated and Sensitive Metabolomics Platform for Human Disease Prediction, Diagnosis and Treatment"
Norman J Dovichi	"Glycolipid Metabolism in Single Cells"
Ronald Breaker	"Sensing Metabolites with Riboswitches"
John D York	"Biological Oscilloscopes: Spatio-Temporal Metabolomics"
Dr. Alan M Kleinfeld	"Profiling unbound metabolites using fluorescent probes"
Dr. James C Liao	"Automated Chip-Based Metabolomics Analysis"
r. Henri Brunengraber	"Dynamic Metabolomics via Isotopomer Analysis"

Awarded Projects (R01)

- Cravatt, Benjamin F (NCI)
 Chemical Probes for Metabolic Pathway Discovery in Human Disease
- Freyer, James P (NCI) Differential metabolic network analysis of tumor progression
- Han, Xianlin (NIA) Shotgun Lipidomics and Alterations in Sphingolipidomes in Alzheimer's Diseases
- Rabinowitz, Joshua D (NAID) Metabolomics of the Virus-host Cell Interaction
- Sweedler, Jonathan* (NIDCR) The Neurometabolome of a Sensory Neuronal Network
- Burant, Charles F (NIDDK) Using Systems Biology to Understand Islet Adaptation and Failure Diabetes
- Frommer, Wolf B* (NIDDK) Sugar signaling networks detected by high content fluxomics
 - bvichi, Norman J*(NINDS) Glycolipid metabolism in single cells

Scientific Progress

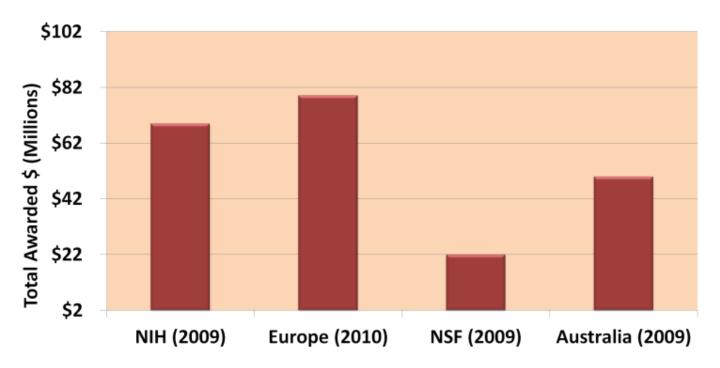
- NCRR meeting Sept. 17th, 2010
 State of Metabolomics Technologies in Translational Research
- Application in clinical/epidemiological studies
- Increasing depth, breadth, throughput
- Quantification
- New Tech (i.e. Metabolomics Imaging)

http://videocast.nih.gov/

Note: BPC3 plans to propose metabolomics as the functional aspect of the project in their renewal application

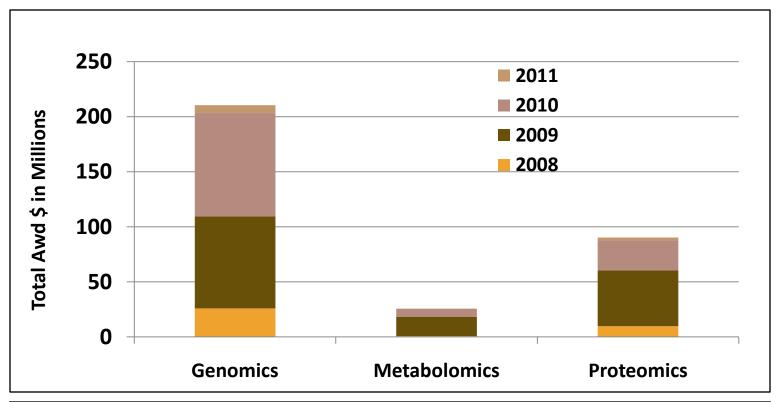
INTERNATIONAL EFFORTS

- Australia Metabolomics Bioplatforms: \$208 million
- Netherlands Metabolomics Center: \$67 million
- BBSRC, UK Plant and Microbial Metabolomics: \$10.4 million
- Canadian Human Metabolome Database: \$8.1 million
- Global investment in Metabolomics for FY 2010 is Estimated to be \$ 225 million*



^{*} Current trends in the Metabolomics sector within Europe. http://www.frost.com/prod/servlet/market-insight-top.pag?docid=128394679

Comparing the NIH Investment in Other "Omics" Fields in Translational Research (FY 2008-2011)

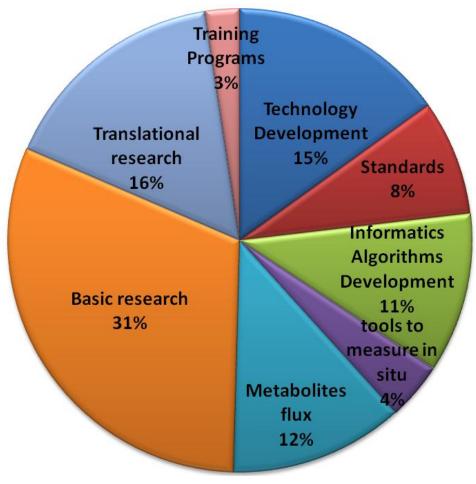


"Omics" Fields	2008	2009	2010	2011	Grand Total
Genomics	\$ 25,913,961	\$ 83,529,488	\$ 93,877,672	\$7,137,180	\$ 210,458,301
Metabolomics	\$ 760,996	\$ 17,620,677	\$ 7,125,183	\$ 149,982	\$ 25,656,838
Proteomics	\$ 9,671,349	\$ 50,798,279	\$ 27,025,316	\$2,756,224	\$ 90,251,168
Grand Total	\$ 36,346,306	\$151,948,444	\$ 128,028,171	\$10,043,386	\$ 326,366,307

NIH Investment in Metabolomics by Topics

FY 2005 – Present Single Project Mechanisms R, DP, F, K, T

Topic	Total Awarded\$		Number of Projects	
Technology Development	\$	65,464,257	45	
Standards	\$	34,815,891	29	
Informatics Algorithms Development	\$	48,303,827	40	
tools to measure in situ	\$	17,374,348	12	
Metabolites flux	\$	52,587,656	49	
Basic research	\$	134,518,374	128	
Translational research	\$	69,844,145	68	
Training Programs	\$	10,484,773	50	



- •Majority of the portfolio is Basic Research, with over \$ 69 million (16%) invested in Translational Research
- •Percentage calculated based on the total amount of \$211M awarded for Single projects (Mechanisms R, DP, F, K, T)

Foreign Efforts



BBSRC, UK Plant and Microbial Metabolomics: \$10.4M



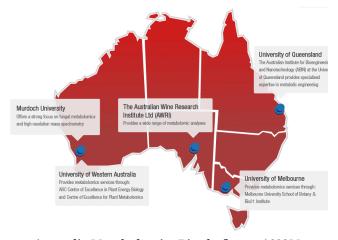
Canadian Human Metabolome Database: \$8.1M



Institute for Advanced Biosciences, Keio University, Japan 100M+

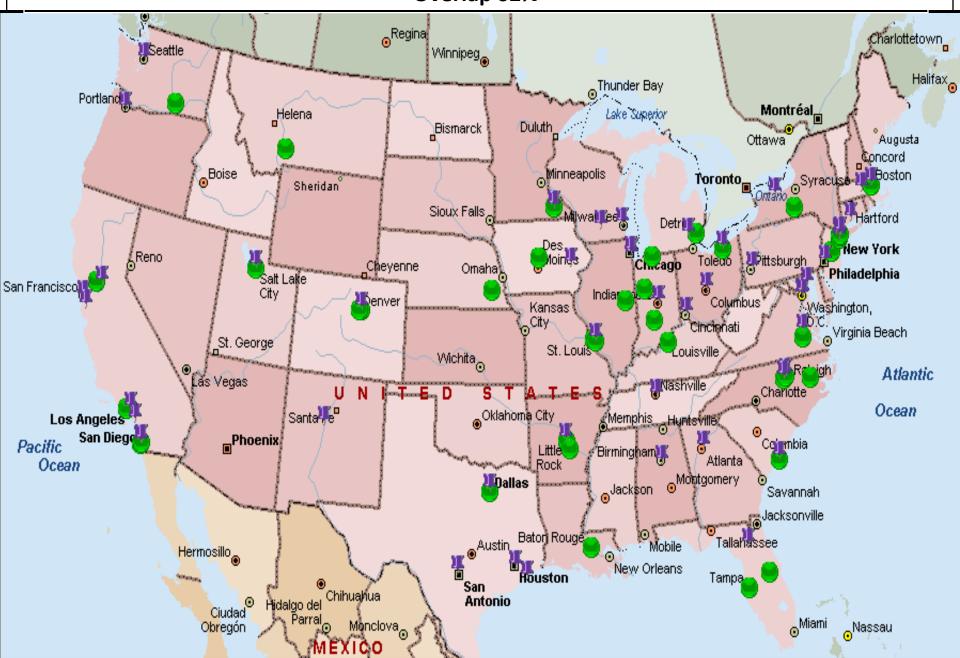


Netherlands Metabolomics Center: \$67M



Australia Metabolomics Bioplatforms: \$208M

Clinical and Translational CTSA Centers (purple) and Metabolomics –Ready Sites(green) Overlap 62%



Common Funds RFAs (\$110 M for 5 years)

RFA-RM-11-016

Regional Comprehensive Metabolomics Resource Cores (RCMRC) (U24)

RFA-RM-11-017

Mentored Research Scientist Award in Metabolomics (K01)

RFA-RM-11-018

Development of Courses or Workshops in Metabolomics (R25)

RFA-RM-11-019

Technology Development to Enable Large Scale Metabolomics Analyses (R01)

RFA-RM-11-020

Metabolomics Data Repository and Coordinating Center (U01)

NOT-RM-11-025

Request for Information on Specific Needs for Metabolomics Reference Standards

NOT-RM-11-024

Notice of Availability for Administrative Supplements for Collaborative Activities to Promote Metabolomics Research

Research Questions (Related to EGRP Mission)

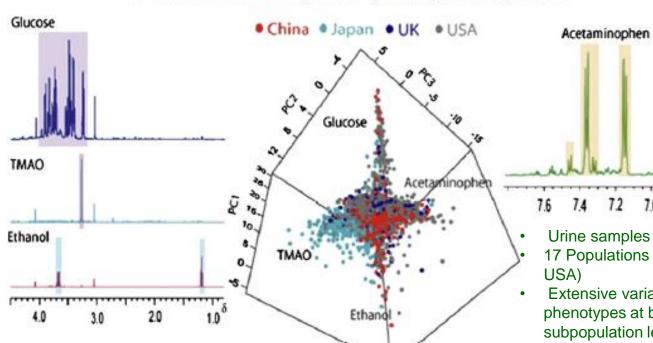
- Can metabolimic <u>profile identify populations at high risk</u> of developing cancer?
- How to <u>integrate information</u> from genomics, epigenomics, proteomics, transcriptomics, microbiome to metabolomics
- Developing metabolomics based <u>mathematical models and</u> <u>algorithms</u> that will be useful for <u>monitoring the progression</u> of cancer and predicting chances for successful treatment
- Probing metabolic pathways of <u>responses to environmental</u> <u>exposures</u> and how these pathways affect <u>individual susceptibility</u> to exposures
- Identifying and validating metabolomics biomarkers (profiles) of <u>biological responses</u> to <u>exposure</u> including compensatory responses and <u>early indicators of cancer</u>
- Develop <u>study design to measure response</u> to environmental changes and disease development by following metabolomic profiles

Research Questions (Related to EGRP Mission)

- Can metabolimic <u>profile identify subtypes of a cancer</u>?
- Can unidentified metabolotes be used for cancer detection and diagnosis (one example in ovarian cancer exists) (common libraries do not have these unidentified metabolites)?
- How can we used metabolomic profiling for personalized medicine?

Large Scale Metabolomic Screening of Human Population

M. Bictash et al. / Journal of Clinical Epidemiology 63 (2010) 970-979

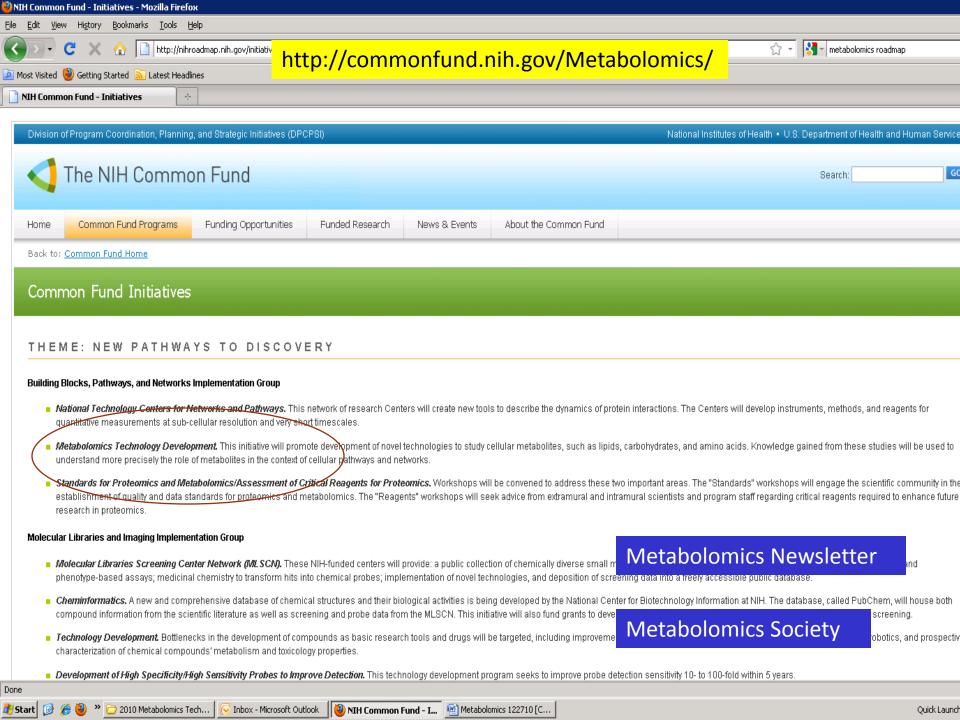


- Urine samples (4,630 people) 17 Populations (China, Japan, UK, USA)
- Extensive variation in metabolomic phenotypes at both country and subpopulation level
- Several biomarkers discriminated significantly between populations

Fig. 1. Large-scale metabonomic screening of human populations. Scores plot for a principal component analysis model derived from ¹H NMR urine data from INTERMAP (n = 4,630 participants), colored by population sample. The ¹H NMR measurement for a given urine specimen is plotted as a single point. The plot shows four outlying phenotypes based on latent information extracted from the spectral data sets, attributable to high urinary glucose, TMAO, ethnol, and acetaminophen. Representative spectra of participants with characteristic spectral profiles for these metabolites are also highlighted. NMR, nuclear nagnetic resonance; PC, principal component; TMAO, trimethylamine-N-oxide.

28

772



NIH COMMON FUNDS METABOLOMICS: RESEARCH PRIORITIES

- 1. Support generation of more metabolite standards through coordinated collaborative approach
- 2. Train more scientists in Metabolomics
- 3. Increase the capacity in Metabolomics through establishing more centers
- 4. Support Metabolomics technology development

Percent of NIH Budget



➤ SBIR: Set-aside program for small business concerns to engage in Federal R&D with the potential for commercialization

Set Aside

2.5%

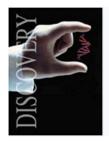
STTR: Set-aside program to facilitate cooperative R&D between small business concerns and U.S. research institutions with potential for commercialization

0.3%

~\$110 million annually at the NCI ~\$650 million annually at the NIH

Small Business Innovation Research (SBIR) Small Business Technology Transfer (STTR)

Programs



PHASE I – Feasibility Study

- Average award: \$170K
- Project period varies, most 6 12 months



PHASE II – Full R&D

- Average \$850K, 2 years but some longer
- Commercialization plan required

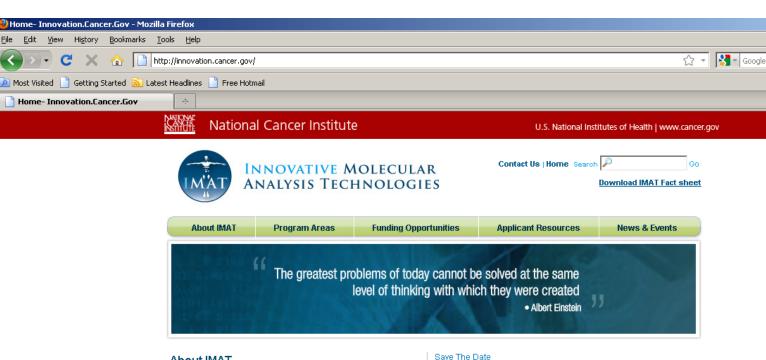


PHASE III – Commercialization

- Use of non-SBIR/STTR Funds
- Consider exit strategy

Issues

- Budgets inadequate for expectations
- Gap between Phases I and II can be almost two years
- Reviewers do not understand challenges of the "D" in "R&D"



About IMAT

The Innovative Molecular Analysis Technologies (IMAT) program was established to support the development, technical maturation, and dissemination of novel and potentially transformative next-generation technologies through an approach of balanced but targeted innovation. In support of its mission, the IMAT program utilizes a variety of investigatorinitiated research project grant mechanisms while retaining a strong commitment to diversity and to the training of scientists and clinicians in cross-cutting, research-enabling disciplines.

Learn More About IMAT +

Recent News

TransGenomic Corporation Accelerates Acquisition of IMAT **Funded Technology**

Transgenomic has announced that it has licensed a high-sensitivity mutation detection technology called Cold-PCR from the Dana-Farber Cancer Institute (DFCI) in Boston, more +

RiboMed Receives \$150,000 Epigenetics Research Contract to Continue Work Funded through IMAT to Develop Better **Detection Technologies for Cancer more +**

Second IMAT investigator named as finalist for ALA Innovation of the Year Award

For the second time in two years, an IMAT investigator has been named as a finalist for the American Laboratory Association's prestigious Innovator of the Year Award, more +

United States Senator featured as Keynote Speaker at IMAT

Current Funding Opportunities

Authorization for the IMAT program expired at the end of calendar year (CY) 2010. The new NCI leadership will be carefully reviewing the IMAT program, and the NCI hopes to reauthorize IMAT in CY2011 and release IMAT funding opportunity announcements (FOAs) in the early part of CY2012. This information will be provided on this website once authorization has been granted.

_ B ×

Please see the links below for past funding opportunities to understand the FOAs we hope to announce with reauthorization of the program.



Funding opportunities in technology development are intended to support the inception, maturation, and subsequent dissemination of technically innovative and potentially transformative emerging

IMAT Supported Commercial Technologies





IMAT Award: Enzymatic Tools for Degrading Tissue and Preserving RNA (R43 – 2001,R44, 2005-2007)

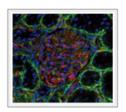
Pl: Gary Latham, Ph.D

Ambion^{*}



IMAT Award: Protein Profiling Arrays, Random Arrays for Gene Expression Profiling (R43 – 1998, R44- 1999)

PI: Mark Chee, Ph.D

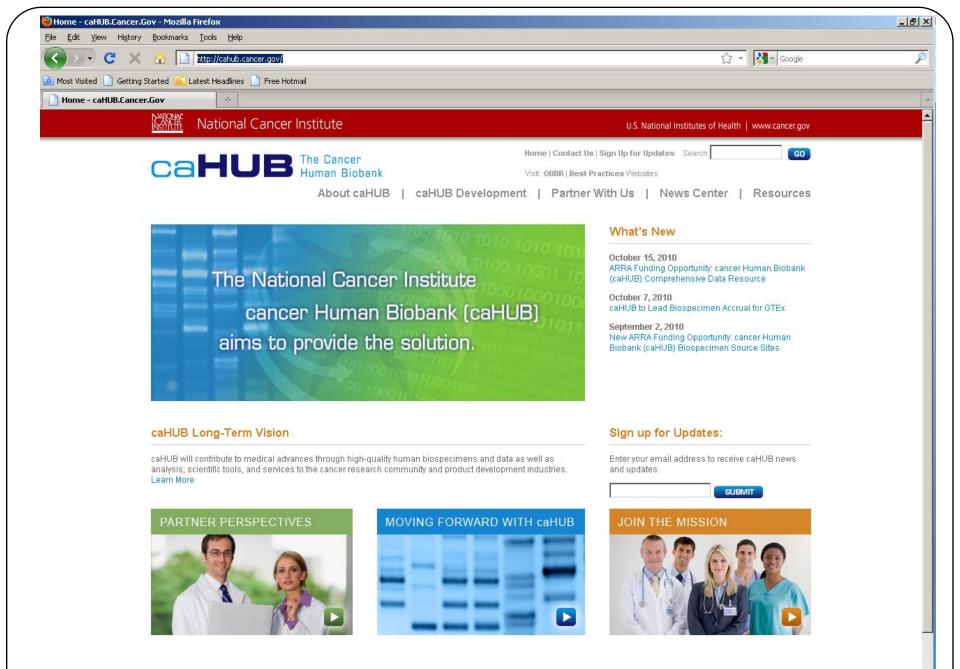


invitrogen

Caı **Quantum D**o

IMAT Award: Sensitive, Multiplexed Analysis of Breast Cancer Markers (R44 - 1999)

Quantum Dot Corp, PI: Robert H. Daniels, Ph.D.,



Biospecimen Lifecycle: Pre-analytical Factors OBBR Office of Biorepositories Affect Molecular Composition and Integrity

⇒Specimen is <u>viable</u> and biologically reactive

Time 0

Molecular composition subject to further alteration/degradation

Factors (examples):

- Antibiotics
- Other drugs
- Type of anesthesia
- Duration of anesthesia
- Arterial clamp time

Factors (examples):

- Time at room temperature
- Temperature of room
- Type of fixative
- Time in fixative
- Rate of freezing
- Size of aliquots



Public-Private Partnership Program Examples





accelerate identification, development and regulator, qualification of biomarkers for: cancer, inflammation and immunity, metabolic disorders, and neuroscience

Alzheimer's Disease Neuroimaging Initiative - NIA, NIBIB, FDA -





Federal partners: NIAMS, NIA, NIDCR, ORWH, NCCAM, NIMHD

Private partners: Merck, Novartis, Pfizer

Publically available database of gene-gene association studies -NIH, Pfizer, Affeymetrix, Abbott -







Grand Challenges

•• Global Health

effective, inexpensive, simple health tools for low resource settings

NIH-FDA Regulatory Science Initiative

- a Common Fund Program

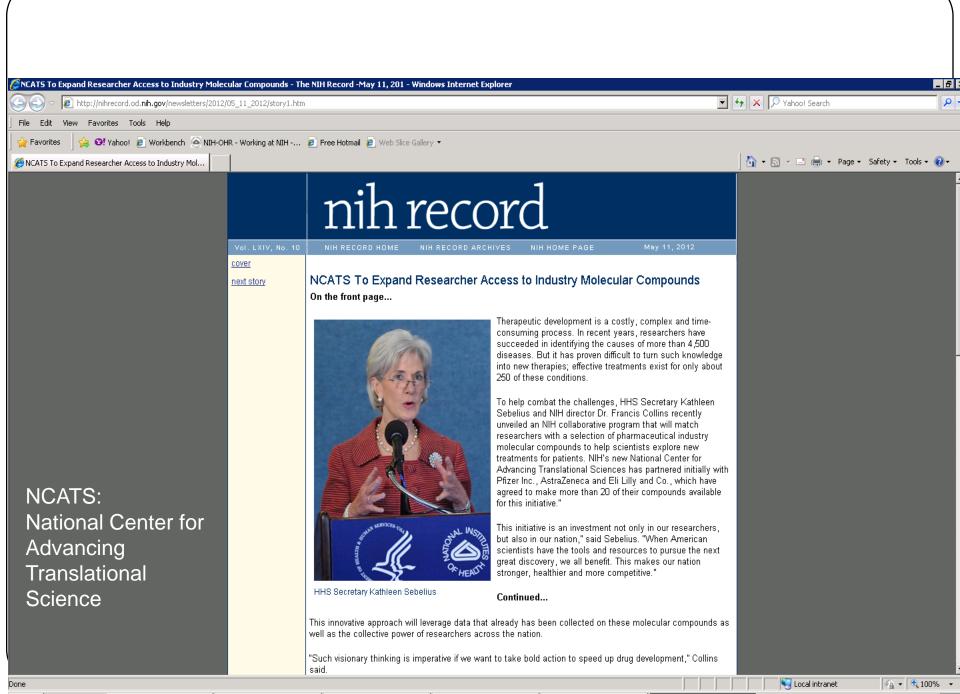
development and use of the scientific knowledge, tools, standards, and approaches necessary for the assessment of medical product safety, efficacy, quality, potency, and performance







- ACCELERATING DRUG AND DEVICE EVALUTATION THROUGH INNOVATIVE CLINICAL TRIAL DESIGN (University of Michigan: William Barsan, Roger Lewis, Donald Berry)
- REPLACEMENT OCCULAR BATTERY (ROBATT) (MB Research Labs: Daniel Cevern, George DeGeorge)
- CHARACTERIZATION/BIOINFORMATICS-MODELING OF NANOPARTICLE:COMPLEMENT INTERACTIONS (University of Washington: Dennis Hourcade)
- HEART-LUNG MICROMACHINE FOR SAFETY AND EFFICACY TESTING (Harvard University: Donald Ingber)

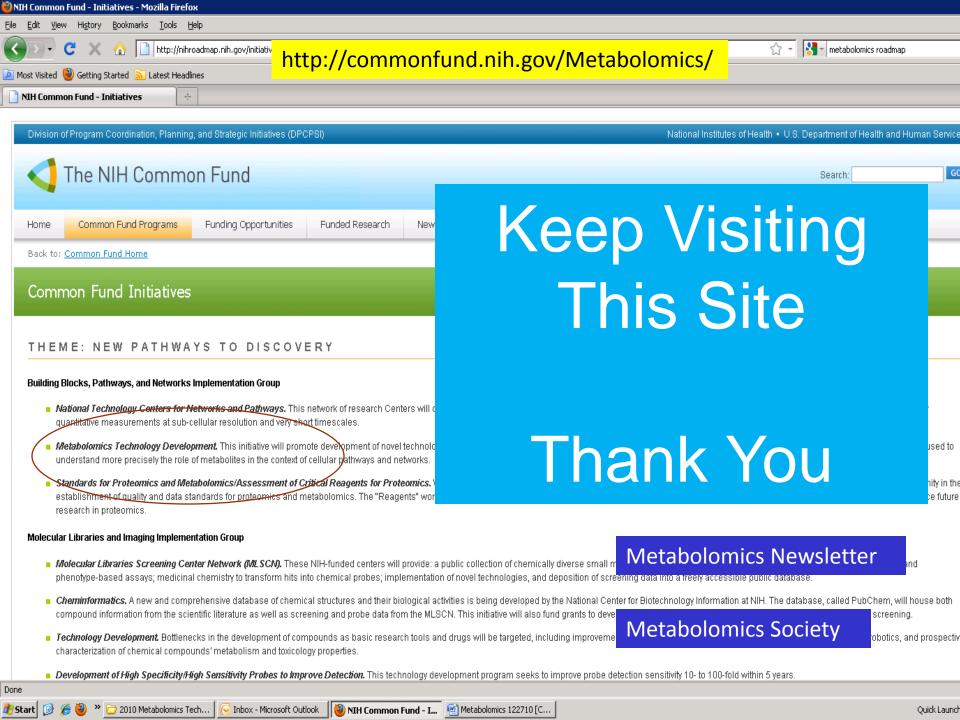




- What is the relationship between low frequency mutations and "driver" mutations? How to determine which mutations have key roles in tumor development?
 - Can we establish methods that will determine which changes are important for tumor development and use these methods to study the functional roles of these mutations?
 - Appropriate mutation analyses will provide an important set of RNA and protein targets for therapy and yield key insights to cancer etiology

Technology opportunity:

- Tools to elucidate the relationship between individual and pathway-related mutations and tumor development
- Tools to track order of mutation development and corresponding metastatic potential in models



Let Us Meet Again

We welcome you all to our future conferences of OMICS International

Please Visit:

www.metabolomicsconference.com

www.conferenceseries.com

http://www.conferenceseries.com/clinical-

research-conferences.php