

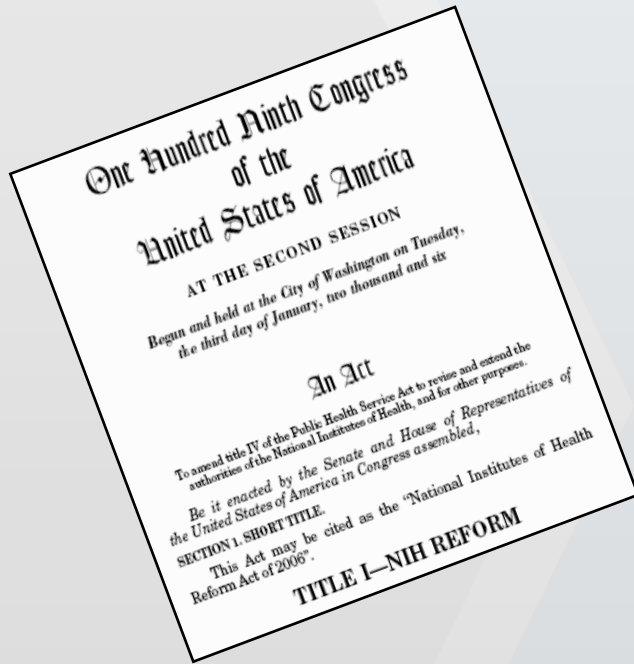
# NIH Common Fund Metabolomics Program

Keren Witkin, PhD

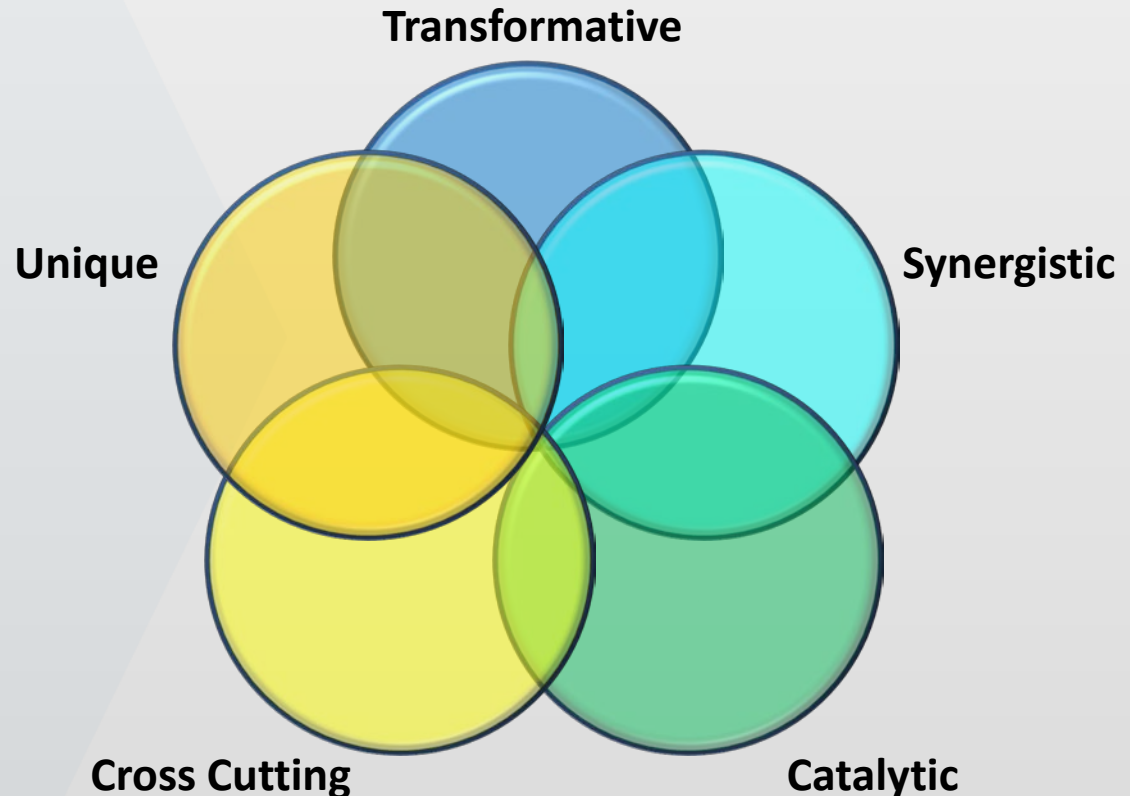
Program Director, CF Metabolomics Program  
Division of Cancer Biology, NCI, NIH

Sept 27, 2016

# The NIH Common Fund: A Different Approach to Science Management



- Exceptionally high impact
- Broadly applicable
- Benefit from strategic planning and coordination

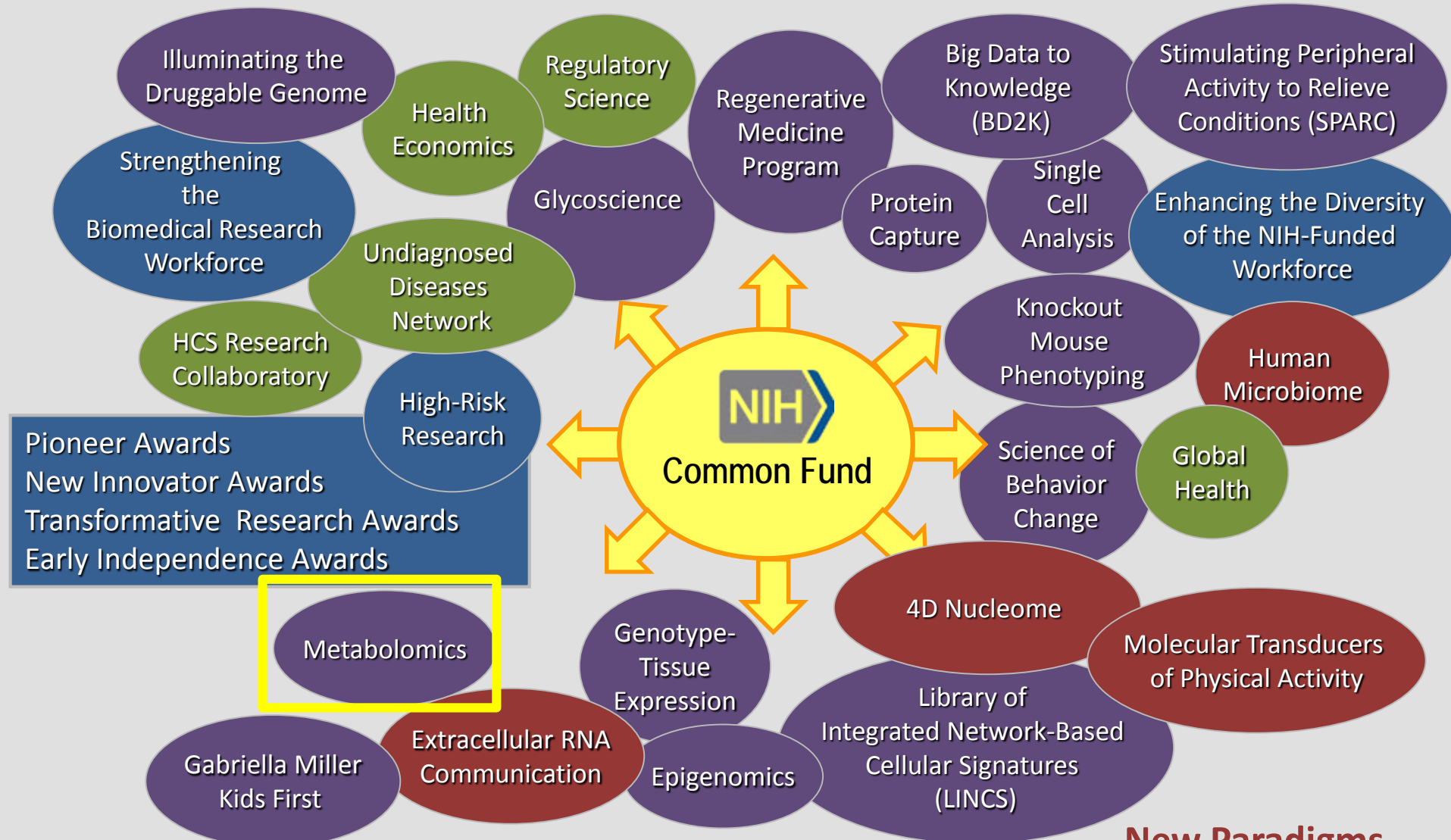


Managed by the Office of Strategic Coordination in the Office of the Director, NIH

# Current Common Fund Programs (FY16)

## New Types of Clinical Partnerships

## Transformative Tools/Methods



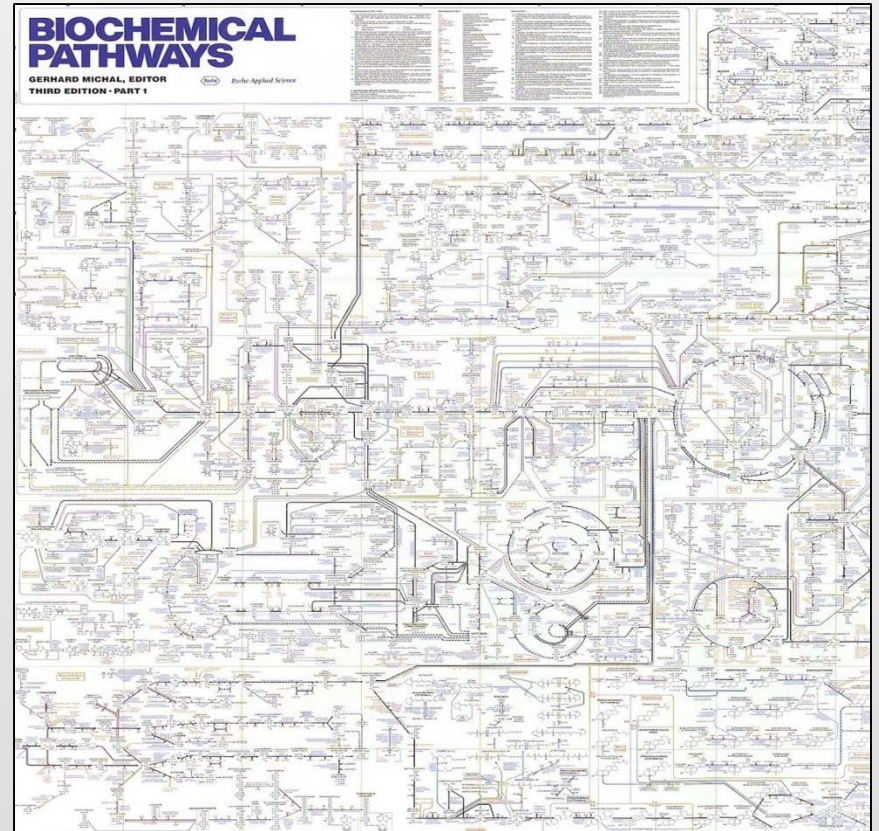
## Transformative Workforce Support

## New Paradigms

# Why Metabolomics?

**Metabolomics:** The systematic study of the metabolites in cells, tissues, and a variety of biospecimens, typically utilizing MS or NMR analytical platforms.

- High potential to contribute greatly to our knowledge of health and disease.
- Applicable to basic, clinical, and translational research.
- Trans-NIH relevance.
- Likely to benefit from strategic coordination.



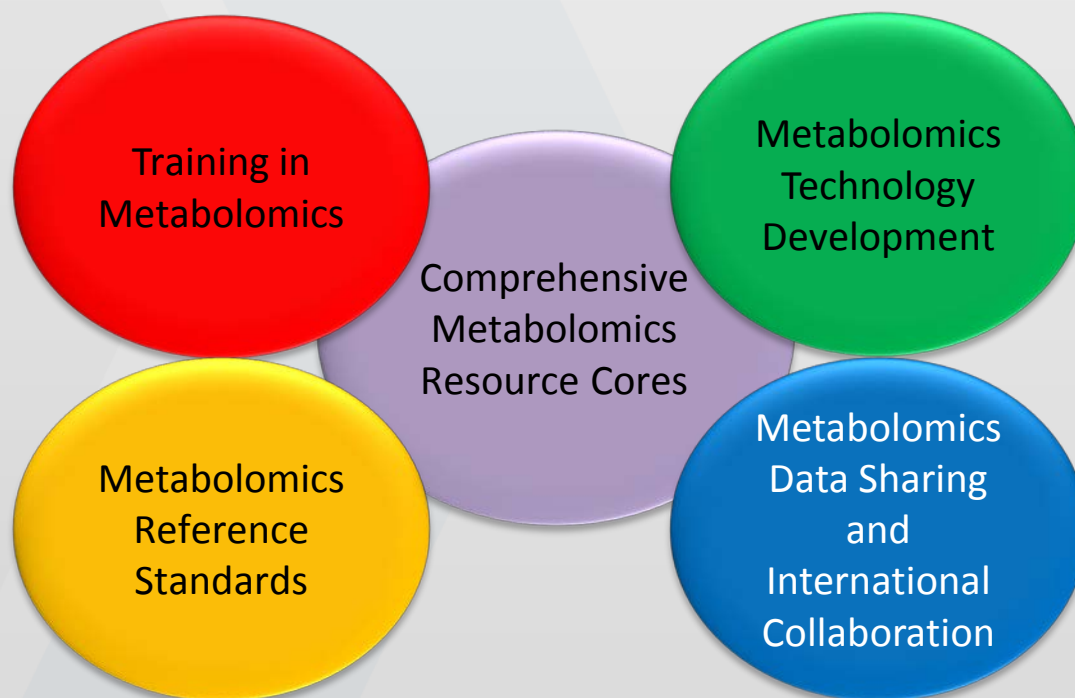
[Biochemical Pathway Map courtesy of Roche](#)

# Assessing Outstanding Needs

- Development of **new technologies** and adoption of existing technologies and methods.
- Availability of **metabolite reference standards** and mechanisms to make new ones as needed.
- **Specialized facilities** that provide high quality metabolomics data, analyses, and interpretation available for collaboration or fee-for-service.
- **Training** for biomedical scientists in the technology, biochemistry, and bioinformatics needed for metabolomics studies.
- A centralized location to store **high-quality metabolomics data** and provide tools for **data analysis**.

# Rationale for the Metabolomics Program

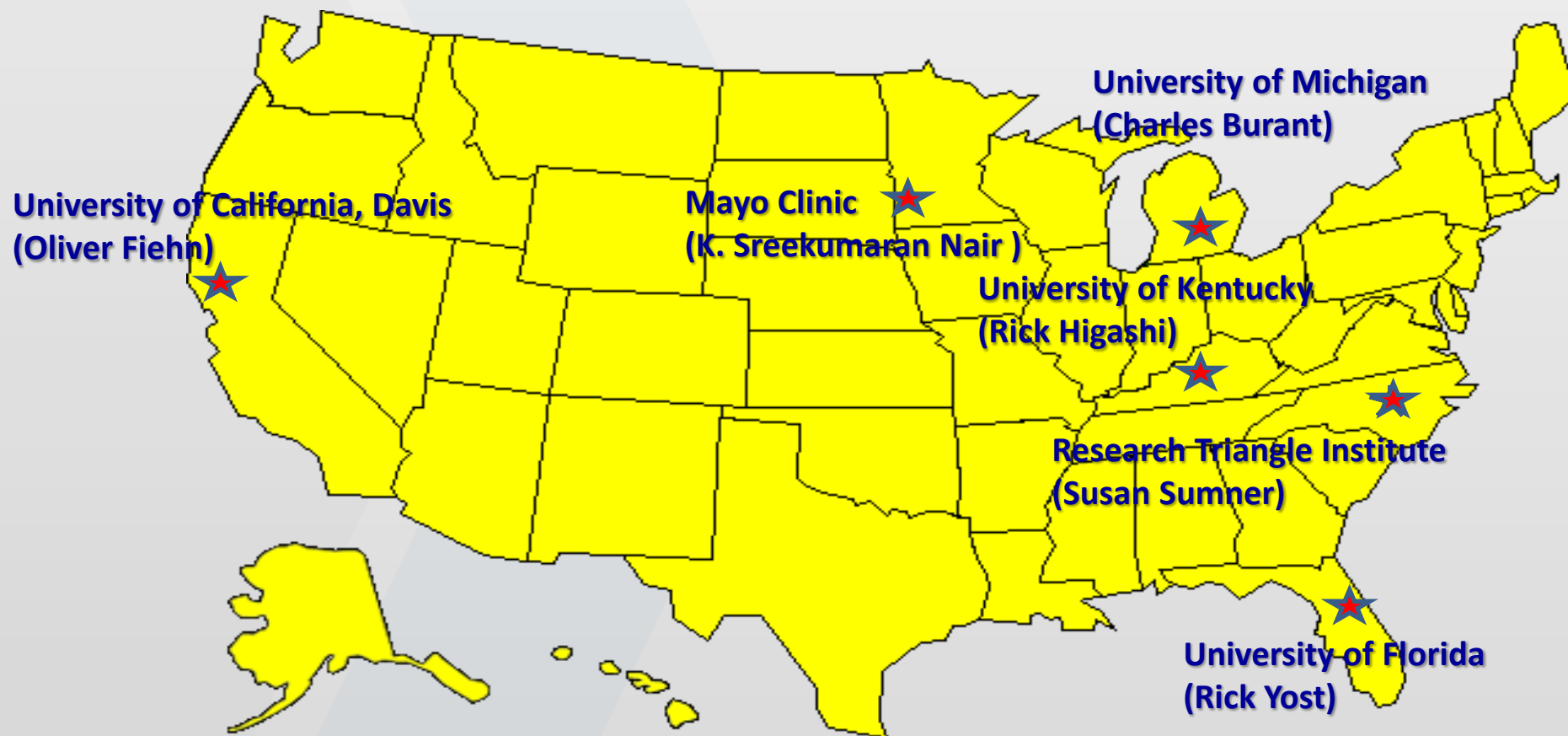
The Common Fund Metabolomics Program was initiated in 2012 **to increase the national capacity in metabolomics** by developing:



## To Meet These Goals:

- Regional Comprehensive Metabolomics Resource Cores (RCMRCs)
- Data Repository and Coordinating Center (DRCC)
- Technology Development (R01) Awards
- Mentored Research Training (K01) Awards
- Grants to Develop Metabolomics Courses
- Metabolite Standards Synthesis Contracts
- Administrative Supplements to Existing NIH Grants
- Pilot and Feasibility (P&F) Awards
- Data Analysis Grants (R03)

# The Regional Comprehensive Metabolomics Resource Cores (RCMRCs)



Visit their individual webpages or access through: <http://www.metabolomicsworkbench.org/>



## Goals of the RCMRCs:

- **Provide metabolomics services for the research community**
  - Fee-for-Service (at cost)
  - Collaborative Pilot and Feasibility (P&F) Program
- **Technology development**
  - Data generation and analysis
- **Metabolomics workforce development**
- **Consortium member - enhancing the field of metabolomics**
- **Data Sharing**
- **Build customer base to become self-sustaining**

# RCMRC Accomplishments:

**159** P&F awards since 2013  
**73** % outside home institution

Workshops

Symposia

**1781** service requests accepted in past 2 years  
**39** unique institutions/year per RCMRC

**289** publications in past 2 years

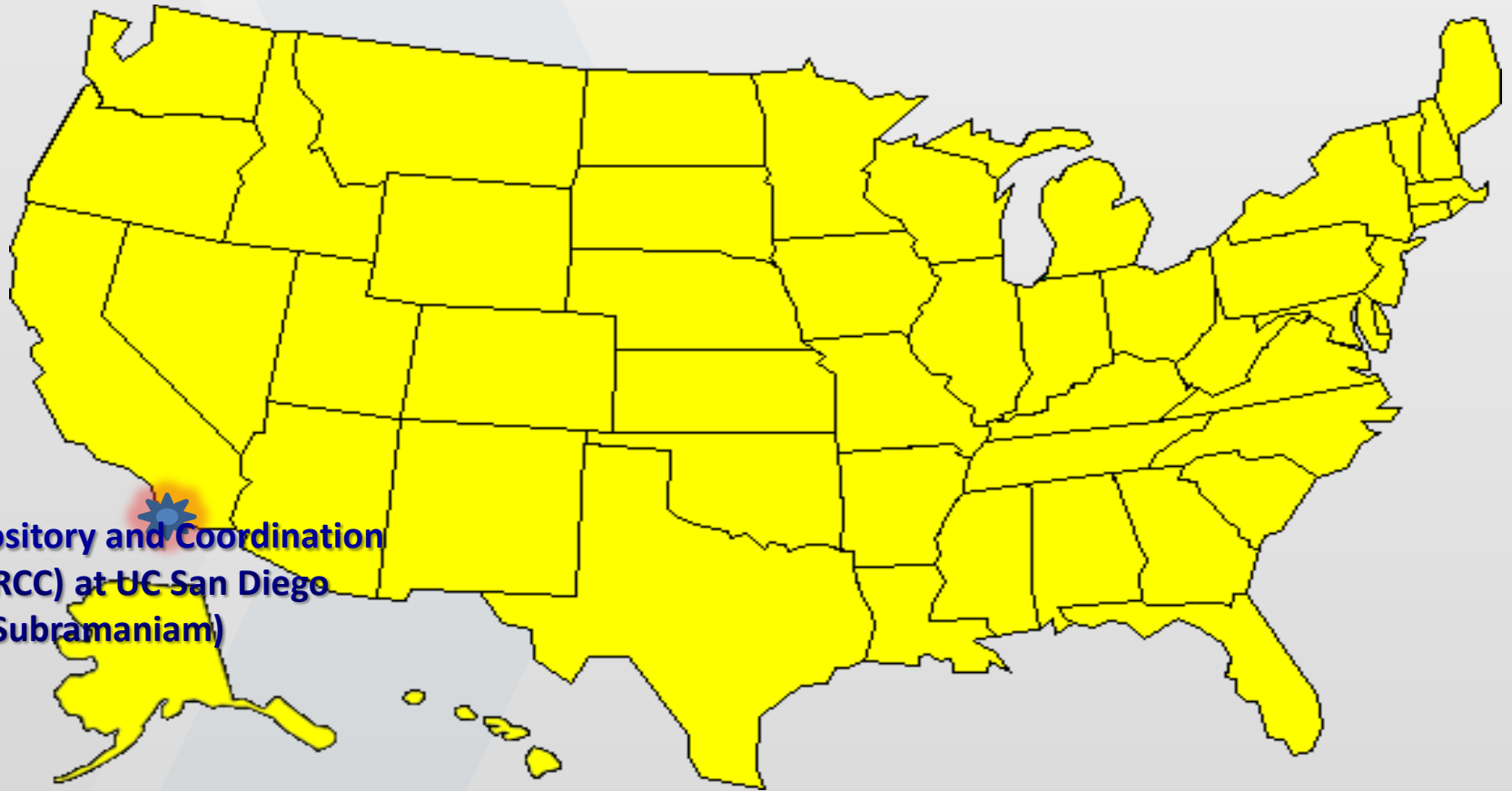
Individual training

**307** datasets deposited

➤ RCMRCs on target to be financially self-sufficient

Data Sharing  
and  
International  
Collaboration

# Data Repository and Coordinating Center (DRCC)



**Data Repository and Coordination  
Center (DRCC) at UC San Diego  
(Shankar Subramaniam)**

Visit their individual webpages or access through: <http://www.metabolomicsworkbench.org/>

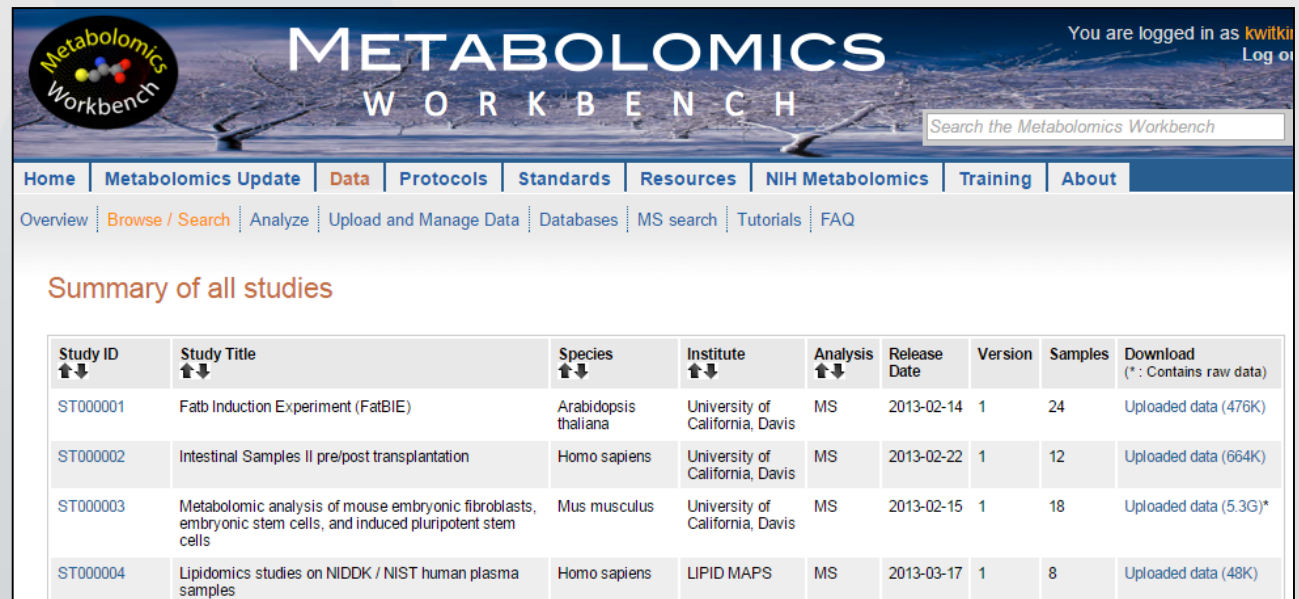
## Goals of the DRCC:

- Develop **data repository** to accept high quality metabolomics datasets from a wide variety of studies
- Collaborate with **international colleagues** to develop minimum requirements for submission of data
- Coordinate **consortium activities** to maximize exchange of best practices and technical advances
- Develop overall **Promotion and Outreach** plan for the Program

# DRCC Accomplishments:

- Developed **data repository** for raw spectra and processed metabolomics data
- Created a **web portal** for consortium activities and resources including datasets, analytical tools, training, protocols etc.

- **492 registered users**
- **371 datasets deposited**



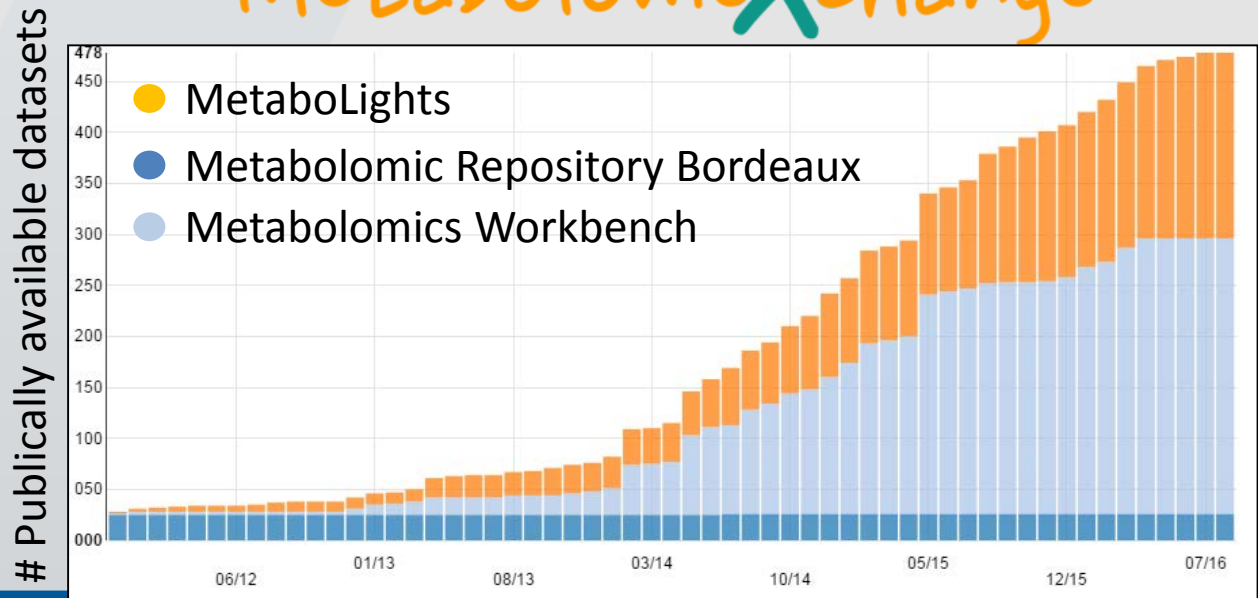
The screenshot shows the Metabolomics Workbench website interface. At the top, there is a navigation bar with links for Home, Metabolomics Update, Data, Protocols, Standards, Resources, NIH Metabolomics, Training, and About. Below this is a search bar and a secondary navigation bar with links for Overview, Browse / Search, Analyze, Upload and Manage Data, Databases, MS search, Tutorials, and FAQ. The main content area displays a "Summary of all studies" table with the following data:

Study ID ↑↓	Study Title ↑↓	Species ↑↓	Institute ↑↓	Analysis ↑↓	Release Date	Version	Samples	Download (* : Contains raw data)
ST000001	Fatb Induction Experiment (FatBIE)	Arabidopsis thaliana	University of California, Davis	MS	2013-02-14	1	24	Uploaded data (476K)
ST000002	Intestinal Samples II pre/post transplantation	Homo sapiens	University of California, Davis	MS	2013-02-22	1	12	Uploaded data (664K)
ST000003	Metabolomic analysis of mouse embryonic fibroblasts, embryonic stem cells, and induced pluripotent stem cells	Mus musculus	University of California, Davis	MS	2013-02-15	1	18	Uploaded data (5.3G)*
ST000004	Lipidomics studies on NIDDK / NIST human plasma samples	Homo sapiens	LIPID MAPS	MS	2013-03-17	1	8	Uploaded data (48K)

# DRCC Collaborative Accomplishments:

- Developed minimal **metadata standards**
- Created a **reference directory** of metabolite names
- Coordinated an inter-lab **reproducibility** exercise
- Worked with international metabolomics community to promote **data sharing**

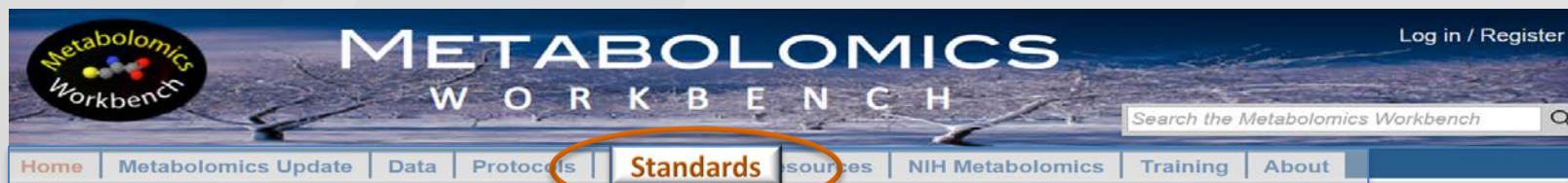
MetabolomeXchange



# Metabolite Standards Synthesis Core

- **High-quality metabolite standards** are synthesized under contracts to SRI International and RTI International
- Provides metabolomics researchers with high quality metabolite standards that have high potential in translational research at no cost
- 20 standards completed; 35 in process

**Nominate a compound or request an aliquot today!**



View the list of nominated compounds or nominate new compounds at:

<http://www.metabolomicsworkbench.org/standards/index.html>

# Metabolomics Technology Development

Project	PI	Technology	Publications
R01ES022181	Patti	Untargeted Workflow	21
R01ES022186	Patterson	Metabolite Extraction Standardization/ Optimization	20
R01ES022191	Fan	Chemoselective Probes	15
R01ES022190	Baker	Ultrarapid Chemical Separation	11
R01ES022176	Hu	Tissue NMR	7
R01ES022172	Murphy	Lipid Identification	3
<b>Total</b>			<b>62</b>

- Program has improved the extraction, separation, detection and identification of metabolites





# Enhancing the Metabolomics Workforce

K01 mentored training  
P&F projects  
Admin Supplements  
Internships/sabbaticals



Small group  
workshops:  
R25 UAB course  
RCMRCs



RCMRC  
symposia



On-line course



Individual  
instruction

Reaching  
the masses



Supported by NIH/NIDDK award R25GM103798 and R25GM103802

# The NIH Investment in Metabolomics Continues to Increase



# Striving to Meet the Evolving Needs of the Metabolomics Community

- Continuing to enhance the **Metabolomics Workbench** in response to community feedback
- Improving the **tools** available for metabolomics studies, including data analysis and interpretation
- Offering **training** at all levels
- Working to identify and promote **best practices** in study design, data acquisition, and data analysis



# Common Fund Metabolomics Working Group

## Chairs:

NCI Dinah S. Singer

NIDDK Philip Smith

## Coordinators:

NCI Barbara Spalholz

NIDDK Arthur L. Castle

OD Leslie Derr

## Project Team Leaders:

NCI Keren Witkin (ME Program Director)

NHLBI Pothur R. Srinivas (Reference Standards)

NIDDK Padma Maruvada (RCMRCs, DRCC)

NIEHS David M. Balshaw (Technologies)

NIGMS Richard T. Okita (Training)

NCATS Danilo Tagle

NCI Mukesh Verma

NCI Krista Zanetti

NHGRI Lita M. Proctor

NHLBI Simhan Danthi

NIA Yih-Woei Fridell

NIAAA Gary J. Murray

NIAID Conrad Malia

NIAMS Hung Tseng

NIDA Amy Lossie

NIDCR Lillian Shum

NIDDK Leeanna Arrowchis

NIEHS Andy Maynard

NIEHS Dan Shaughnessy

NIMH Laurie S. Nadler

NINDS Katrina Gwinn

OD Aron Marquitz

OD Nicole Lim