

OCTOBER 1, 2009

...To Meet Your  
Research Needs  
In Diabetes,  
Endocrinology,  
and Diabetes  
Complications...

**NEW WEBSITE**  
<http://DERC.ucsd.edu>

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Please remember to cite the DERC Grant in all papers that utilize DERC Cores or are supported by the Pilot and Feasibility Awards:

"Our research utilized Core (or Research) support from the UCSD/UCLA NIDDK Diabetes and Endocrinology Research Center P30 DK063491."

## THREE EXCITING FALL MEETINGS FOR UCSD/UCLA/SALK/CEDARS DERC MEMBERS:

**METABOLISM: BENCH TO BEDSIDE, OCT. 8-10, LA JOLLA**  
**WESTERN JOINT DERC MEETING, OCT. 10, LA JOLLA**  
**USC LIVER CENTER AND DERC JOINT MEETING,  
DEC. 4, LOS ANGELES**

*DETAILS INSIDE!*

## ARRA FUNDS AWARDED TO THE DERC!

ARRA Funds were awarded to our DERC in the amount of \$500,000 direct costs over two years. The funds are specifically to DOUBLE our P&F grants for 2009 and 2010 to \$300,000 per year and to purchase large equipment for the Core Facilities.

## 2009 DERC P&F Grants Awarded Pilot and Feasibility Projects in Endocrinology and Diabetes

Pilot & Feasibility Program, Director: Pinchas Cohen

As part of the ARRA Funds awarded to our UCSD/UCLA DERC grant, the Pilot and Feasibility grant program will support 8 grantees at approximately \$30,000-\$40,000 per year for 2009 and 2010, double the number normally available.

Thus, the upcoming 2010 competition will award \$300,000 in awards for P&F. Watch for the announcement for the competition early in 2010!

### THE UCSD/UCLA DERC is Proud to Announce the 2009 P&F AWARDEES:

#### **New ARRA P&F Awardees for 2009:**

**Gregory Harmon**, Clinical Instructor, Department of Medicine, Division of Gastroenterology, UCSD

*Peroxisome Proliferator-Activated Receptor Gamma Ameliorates the Phenotype of Cystic Fibrosis*

**Michael Downes**, Senior Staff Scientist, Gene Expression Laboratory, The Salk Institute for Biological Sciences, La Jolla

*Deciphering Nuclear Receptor Regulation of Diabetes via Promoter Ontology analysis*

**Jamie Powers**, Clinical Instructor, Mattel Children's Hospital, Department of Pediatrics, UCLA

*Effects of Hyperglycemia on Adrenal Cortex Function and Steroidogenesis*

**Senta Giorgia**, Assistant Adjunct Professor, Endocrinology, Diabetes, and Hypertension, Department of Medicine, Geffen School of Medicine, UCLA  
*Using Comparative Methylation and Gene Expression Analysis to Understand Age-Restricted Beta Cell Replication in Response to Insulin Resistance*

#### **Original 2009 awardees:**

**Mina Desai**, Associate Professor, UCLA

*Development of Insulin Resistance and Adiposity via PPAR Dysregulation*

**Anthony Heany**, Associate Professor, UCLA

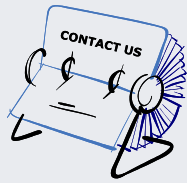
*Role of GLUT5 in Pathogenesis of Metabolic Syndrome*

**Karen Herbst**, Assistant Professor, UCSD

*Blockade of Receptor Cleavage in Diabetes Mellitus with an MMP Inhibitor*

**Andrea Hevener**, Assistant Professor, UCLA

*The Impact of Myeloid-Specific ERα Expression on Inflammation, Insulin Action and Adiposity*



#### Listserv for DERC Members

Send announcements, communications, requests, etc., to your DERC colleagues:

#### DERC-L@UCSD.EDU

If you are receiving this newsletter directly, you are already subscribed. If you would like to subscribe, please email mellonadmin@ucsd.edu. This is a moderated listserv, so messages will be prescreened such that only relevant and important messages will reach you.

#### NEW WEBSITE

<http://DERC.UCSD.EDU>

#### Contact information for DERC Cores and Programs:

##### DERC PI/Director:

**Jerrold Olefsky**  
(858) 534-6651  
jolefsky@ucsd.edu

##### Administration:

**Betsy Hansen**  
(858) 534-6651  
ejhansen@ucsd.edu

##### Pilot & Feasibility Program

**Pinchas Cohen, M.D.**  
(310) 206-5844  
hassy@mednet.ucla.edu

##### Transgenic and Knockout Mouse CORE:

<http://cancer.ucsd.edu/tgm/>  
**Pamela Mellon, Ph.D.**  
Core Director

##### Core Contacts:

**Jun Zhao**  
Transgenic Mice Contact  
858-822-3270  
tg@ad.ucsd.edu

##### Ella Kothari

Gene Targeting (Embryonic Stem Cells and Blastocyst Injection) Contact  
858-534-3178  
stemc@ad.ucsd.edu

##### Heather Oakley

Embryo Cryopreservation  
858-822-2108  
cryo@ad.ucsd.edu

## DON'T MISS THE METABOLISM MEETING NEXT WEEK!

### Clinical Investigation Institute/Nature Medicine Bench to Bedside: Metabolism

October 8-10, 2009

<http://www.nature.com/natureconferences/ctri2009/index.html>

#### Special Pricing for DERC P & F Faculty and UC faculty

As part of our Enrichment activities, the DERC will help host a scientific meeting co-sponsored by Nature Medicine and UCSD next Fall in La Jolla. This Nature Medicine meeting is focused on Diabetes and Metabolism and begins at 4 pm the evening of October 8<sup>th</sup>, ending at Noon at Saturday October 10<sup>th</sup>.

For more information visit <http://cme.ucsd.edu/b2b2009>

Speakers Include:

#### Keynote Speaker: Michael Brown (U Texas)

**Helen Hobbs** (U Texas)  
**Gokhan Hotamisligil** (Harvard)  
**Peter Libby** (Harvard)  
**Michael Karin** (UCSD)  
**Paresh Dandona** (SUNY Buffalo)  
**Gerry Shulman** (Yale)  
**Ira Goldberg** (Columbia)  
**Phil Scherer** (U Texas)  
**Barbara Kahn** (Harvard)  
**Chris Newgard** (Duke)  
**Tony Lam** (Toronto)  
**Zofia Zukowska** (Georgetown)  
**Daniel Drucker** (Toronto)  
**David Cummings** (U Wash)  
**Steve Shoelson** (Joslin Diabetes Center)  
**Francesco Rubino** (Cornell Medical Center-NYC)



#### Hilton La Jolla Torrey Pines

10950 North Torrey Pines Road, La Jolla, California

[http://www1.hilton.com/en\\_US/hi/hotel/SANTPHH-Hilton-La-Jolla-Torrey-Pines-California/index.do](http://www1.hilton.com/en_US/hi/hotel/SANTPHH-Hilton-La-Jolla-Torrey-Pines-California/index.do)

**Mouse Phenotyping CORE:****Rajendra Tangirala, PhD**

Core Director

**Pinchas Cohen, MD**

Core Co-Director

**Andrea Hevener, PhD**

Core Co-Director

**David Hwang, PhD**

Core Co-Director

**Core Contacts:****Binrong Liu**

310-825-8499

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**Diana Becerra/Jason Kim**

310-794-6612

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**Transcriptional Genomics CORE:**<http://www.microarrays.ucsd.edu>**Chris Glass, Ph.D.**

Core Director

**Gary Hardiman, Ph.D.**

Core Co-Director

BIOGEM Core Facility

**Nicholas Webster, Ph.D.**

Core Co-Director

VA Genechip Core

**Core Contacts:****BIOGEM Core**

Agilent and Illumina Arrays

Solexa Sequencing

**Colleen Ludka**

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**Genechip Core**

Affymetrix Arrays

454 Sequencing

**Jorge Valencia**

(858) 552-8585 x7100

genechip@vapop.ucsd.edu

**Human Genetics CORE:****Jerome Rotter, M.D.**

Core Director

**Leslie Raffel, M.D.**

Core Co-Director

**Xiuqing Guo, Ph.D.**

Core Co-Director

**Kent D. Taylor, Ph.D.**

Core Co-Director

**Core Contact:****Cynthia Hernandez, R.N.**

Core Contact

(310) 423-1457

Cynthia.Hernandez@cshs.org

**Inflammation CORE:****Peter Tontonoz, M.D., Ph.D.**

Core Director

**Rajendra Tangirala, PhD**

Core Co-Director

**Core Contact:****Rima Boyadjian**

310-206-4622

rboyadjian@mednet.ucla.edu

**PLEASE JOIN US FOR:  
The First Western DERC Meeting**

**October 10th, 2009, from 1:30-6:30 PM**

*Immediately following the Nature Medicine Meeting*

**Grand Ballroom, Salon D, Hilton La Jolla Torrey Pines**

The Western region DERCs (Baylor, UCSD/UCLA, University of Colorado, and University of Washington) have organized into a subgroup and through the Regional DERC Director's committee, we have arranged for P&F recipients from the other Western Centers to attend and present at this meeting.

**No Registration Fee**

*Speakers should load their presentations between 11:00 AM and 1:00 PM  
12:45 PM Box lunches will be available in the foyer*

1:30-1:40 Opening remarks: Jerry Olefsky, Director of the DERC

1:40-2:20 **Keynote Speaker: Professor Ron Evans, The Salk Institute**

2:20-3:30 Introduction to the WDERC: Hassy Cohen, DERC Co-Director

2:30-3:45 **Selected talks: Insulin Action & T2DM**

Michael Downes, UCSD

*Promoter Ontology Analysis of Nuclear Receptors*

Greg Morton, Univ. Washington

*Hypothalamic Signaling and Insulin Sensitivity*

Laura Cobb, UCLA

*Novel Mitochondrial Peptides and their Role in Metabolism*

David Maahs, Univ. Colorado, Denver

*Lipoprotein Sub-fractions and Atherogenesis in Diabetes*

Andrea Hevener, UCLA

*ERα Effects on Inflammation and Insulin Action*

3:45-4:05 **Recent highlights of the UCSD/UCLA DERC Cores**

Peter Tontonoz, UCLA, The Inflammation Core, Core E

*Harnessing Inflammation Markers for Diabetes Research*

Gary Hardiman, UCSD, The Transcriptional Genomics Core, Core C

*New Frontiers in Sequencing*

4:05-4:15 Tea/Coffee break

4:15-5:30 **Selected talks on Islet Biology and T1DM**

Chris Hampe, Univ. Washington

*Anti-GAD Anti-idiotypic Antibodies in T1DM*

Janet Wenzlau, Univ. Colorado, Denver

*The Humoral Autoimmune Response to ZnT*

Ellen Lumpkin, Baylor

*Sensory Innervation and Pancreatic Function*

Vijay Yechoor, Baylor

*Auto-immunity Evading Neo-islets for Diabetes*

Steven Chessler, UCSD

*Neuroigin-Neurexin Interactions in Islet Function*

5:30-6:30 Wine & Cheese and poster-session reception on patio

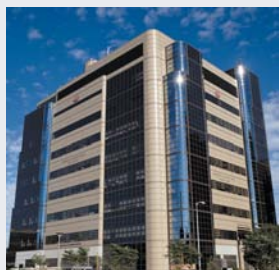
*Poster Boards available from 11:00 AM in the Grand Ballroom*

*Please note that WE ARE STILL ACCEPTING POSTERS FOR THIS MEETING*

*If you wish to present a poster, please email [hassy@mednet.ucla.edu](mailto:hassy@mednet.ucla.edu)*

Key new Publications  
by DERC Members:

Goodarzi, M.O., Taylor, K.D., Jones, M.R., Fang, B., Guo, X., Xiang, A., Buchanan, T.A., Hodis, H.N., Raffel, L.J., Rotter, J.I. 2009 Replication of Calpain-10 genetic association with carotid intima-media thickness, Atherosclerosis, 205:503-505



Ghislette, S., W. Huang, K. Jepsen, C. Benner, G. Hardiman, M.G. Rosenfeld, and C.K. Glass. 2009. Cooperative NCoR/SMRT interactions establish a corepressor-based strategy for integration of inflammatory and anti-inflammatory signaling pathways. *Genes & Development* 23:681-693.



Liu, Y., R. Dentin, D. Chen, S. Hedrick, K. Ravnskjaer, S. Schenk, J. Milne, D.J. Meyers, P. Cole, J. Yates, J. Olefsky, L. Guarent, and M. Montminy. 2008. A fasting-inducible acetylation/deacetylation switch modulates hepatic gluconeogenesis via the CREB coactivator TORC2. *Nature* 456(13):269-273.



## USC Southern California Research Center for ALPD and Cirrhosis JOINT SYMPOSIUM with the UCSD/UCLA DERC

**DECEMBER 4, 2009 at USC in Los Angeles**

8:30-9:00 Continental Breakfast  
9:00-9:10 Welcoming Remarks

### **Mechanisms of Liver Metabolism and Fatty Liver**

9:10-9:35 Barry M. Forman, Ph.D., Professor and Director, Gene Regulation & Drug Discovery, The Beckman Research Inst., City of Hope Med Center  
*"FXR and Hepatic Metabolism: on the Fast Track"*

9:40-10:05 Marc Montminy, Ph.D.  
Professor, Clayton Foundation, The Salk Institute  
*"Regulation of Hepatic Gluconeogenesis by the TORC/CRTC Family of CREB Coactivators"*

10:10-10:35 Mark Czaja, M.D.  
Professor of Medicine, Albert Einstein College of Medicine  
*"Regulation of Cellular Lipid Accumulation by Macroautophagy"*

10:40-10:45 Break

10:45-11:05 Kuk-Wha Lee, M.D., Ph.D.  
Assistant Professor of Pediatrics, UCLA  
*"Contribution of the Growth Hormone/Insulin-Like Growth Factor Axis in the Pathophysiology of Hepatic Steatosis"*

### **Pilot Projects**

11:10-11:25 Ekihiro Seki, M.D., Ph.D., UCSD  
*"TLR Signaling in ASH vs. Non-ASH"*

11:30-11:45 Kinji Asahina, Ph.D., USC  
*"Hepatic Stellate Cell Precursors in Developing and Fibrotic Livers"*

11:50-12:05 Jenny Yuan, Ph.D. UCLA/West LA VA  
*"The Role of Protein Kinase D in Alcoholic Pancreatitis"*

12:10-12:25 Bernd Schnabl, M.D., UCSD  
*"Early Bacterial Translocation in Alcoholic Liver Injury"*

12:30-13:30 Lunch

### **Postdoc Presentations**

13:30-14:30

### **Oxidant Stress, Inflammation, and Cancer**

14:30-14:55 David Brenner, M.D.  
Vice Chancellor for Health Sciences, Dean, UCSD School of Medicine  
*"ROS, NADPH oxidase, and liver fibrosis"*

15:00-15:25 Neil Kaplowitz, M.D.  
Professor and Director, USC Research Center of Liver Disease  
*"Mitochondrial and ER stress Coupling"*

15:30-15:55 Hide Tsukamoto, DVM, PhD  
Professor and Director, ALPD and Cirrhosis Research Center  
*"NASH Models"*

16:00-16:20 Keigo Machida, Ph.D.  
Assistant Professor, Dept of Mol. Microbiology and Immunology, USC  
*"Liver Cancer Stem Cells Generated by HCV, Alcohol, and Obesity"*

16:25-16:40 Break

16:40-17:05 Jerome Rotter, M.D.  
Professor, Pediatrics and Human Genetics, UCLA/Cedars Sinai  
*"Genome-Wide Association of IBD -- From Susceptibility to Therapy"*

17:10-17:35 Anna Gukovskaya, Ph.D.  
Professor and Senior Career Scientist, UCLA, West LA VA  
*"Autophagy in Pancreatitis"*

17:40-18:00 Simon Beaven, M.D., Instructor, Gastroenterology, UCLA  
*"LXR Signaling in Hepatic Inflammation and Fibrosis"*

18:05-18:15 Closing Remarks

18:30-20:30 Reception and Dinner



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<http://DERC.ucsd.edu>



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### **CHECK OUT OUR MEMBERSHIP DIRECTORY**

It allows easy searching by name, keywords, or location, and ONE-CLICK EMAILING to all of your DERC colleagues.



### **CHECK OUT OUR CORES' WEB PAGES**

Find services offered by each Core with leadership and contact information to facilitate your core usage.

### **CHECK OUT OUR ANNOUNCEMENTS PAGES**

See meeting schedules and announcements and one-click connections to the meeting sites and information.



### **CHECK OUT THE PILOT AND FEASIBILITY PROGRAM PAGES**

Apply for P&F grants from the DERC. See the past awardees and get application instructions.

### **CHECK OUT OUR RESEARCH PAGES**

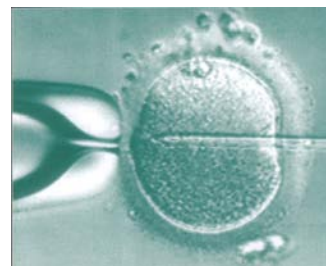
Find important papers by your DERC colleagues and read summaries of their recent work.

## **ARRA EQUIPMENT FUNDS** **\$200,000 direct costs awarded to:**

### **The Transgenic and Knock-Out Mouse CORE** **CORE A. Director: Pamela L. Mellon, Ph.D.**

The DERC Transgenic and Knock-out Core is a state-of-the-art facility that has an outstanding track record in the production of genetically altered subjects. Transgenic subjects carrying new or novel genes are created by microinjection of DNA into the pronuclei of fertilized eggs. Knock-out mice lacking specific genes of interest are created by homologous recombination in embryonic stem cells followed by injection into blastocysts to create chimeric subjects. Highly experienced personnel produce transgenic and knock-out mice for UCSD investigators at subsidized cost and with very short lead times. The Core provides embryonic stem cell recombination, knockout mice, transgenic mice (both standard and BAC transgenics), embryo freezing, and pathogen-free embryonic rederivation to the DERC community at discounted rates. This UCSD-based Core Facility has been in operation since 1992.

**ARRA FUNDS will purchase a third microinjection station which will be equipped for Intracytoplasmic Sperm Injection (ICSI) to allow the testing and development of a service for reconstituting cryopreserved sperm.** Cryopreservation of sperm is a much more cost-effective and straightforward method for preservation of valuable mouse lines allowing DERC members to store their genetically modified mouse lines safely, cheaply and rapidly.



### **The Human Genetics CORE** **CORE D. Director: Jerome I. Rotter, M.D.**

With the rapid advances that have been occurring in the HapMap, high throughput genotyping, and statistical analysis methodology, meaningful advances in the identification of genes contributing to complex disorders such as diabetes have been made. While this progress is exciting, the consensus is that only a small portion of the genes involved in these diseases have been identified and much more work remains to be done. The DERC Human Genetics Core offers expertise to DERC investigators conducting studies into the genetics of diabetes, its complications and related endocrine disorders. The Core provides services necessary for such genetic research in human populations; personnel offer expertise in study design, molecular genetics and statistical genetic analysis and Core facilities are available for high throughput genotyping and establishment of lymphoblastoid cell lines.



**ARRA FUNDS will purchase IBM Blade Center Components and Server and a bay of 15 300GB disk drives.** Large-scale genome wide association study (GWAS) has become a widely accepted approach for locating genes implicated in disease predisposition. In most GWAS studies, 300K to 1 million SNPs are genotyped, resulting in large amounts of genotyping data on each subject. In addition, imputation methods are now commonly used to infer additional genotypes, resulting in as many as 3 million SNPs per individual. The storage of this massive amount of GWAS data, as well as genetic analysis of such data, requires tremendous computing resources. The ARRA-funded computing equipment will increase the rate of data analyses by the Core many fold and accelerate and expand the analysis of results for DERC members.