CARDIO-METABOLIC DISEASE & OBESITY INITIATIVE Faculty Profiles





CENTER FOR METABOLISM AND OBESITY RESEARCH (CMOR)







Michael Wolfgang, Ph.D Director

Dipali Sharma, MS, PhD



Oncology Metabolomics, cancer-obesity connection

Guang William Wong, PhD



Physiology Adipose/muscle hormones, insulin resistance

Todd Tarquin Brown, MD, PhD



Endocrinology Diabetes, Lipid disorders

Peter Espenshade, PhD



Cell biology Cholesterol lipid biosynthesis



Adrian Sandra Dobs, MD, MHS





Endocrinology Diabetes, hyperlipidemia, gonad dysfunction

Jeffery Hunter Young, MD, MHS



Epidemiology Insulin resistance, hypertension, CVD

Lili Ayala Barouch, MD



Cardiology Leptin, b3-adrenergic signaling

METABOLISM AND OBESITY MICHAEL J. WOLFGANG, PHD



Faculty Webpage **Publications Grant Information** **APPOINTMENTS**

Professor, Biological Chemistry Director, Center for Metabolism and Obesity Research

RESEARCH INTERESTS

- nutritional/metabolic cues

RESEARCH **HIGHLIGHTS**

- ketogenic diet
- <u>ketogenesis</u>
- lipid homeostasis
- skeletal muscle





- Understanding molecular mechanisms to sense and respond to

- Exploring novel neuron-specific enzyme function in metabolic processes

- Requirement of hepatic pyruvate carboxylase during fasting, high fat, and

- mTORC1 activation is not sufficient to suppress hepatic PPARα signaling or

- The role of ethanolamine phosphate phospholyase in regulation of astrocyte

- Remodeling glycerophospholipids affects obesity-related insulin signaling in

OBESITY, METABOLISM AND CANCER DIPALI SHARMA, MS, PHD



APPOINTMENTS

Professor, Oncology

RESEARCH INTERESTS

- Understanding the molecular connections between breast cancer and obesity, racial disparities, and microbial dysbiosis
- Developing strategies to abrogate the obesity-breast cancer axis and improving therapeutic responses in obese women

Faculty Webpage **Publications Grant Information**

RESEARCH **HIGHLIGHTS**

- Gut colonization with an obesity-associated enteropathogenic microbe modulates the premetastatic niches to promote breast cancer lung and liver metastasis
- Weight Gain after Hormone Receptor-Positive Breast Cancer
- Hyperleptinemia in obese state renders luminal breast cancers refractory to tamoxifen by coordinating a crosstalk between Med1, miR205 and ErbB
- Concomitant activation of GLI1 and Notch1 contributes to racial disparity of human triple negative breast cancer progression





Member, Center for Metabolism and Obesity Research

METABOLIC DISEASES AND OBESITY **GUANG W. WONG, PHD**



Faculty Webpage **Publications Research Grants**

APPOINTMENTS

Professor, Physiology Member, Center for Metabolism and Obesity research

RESEARCH INTERESTS

- metabolism by circulating factors
- C1q/TNF protein family, PRADC1, CTRP13

RESEARCH **HIGHLIGHTS**

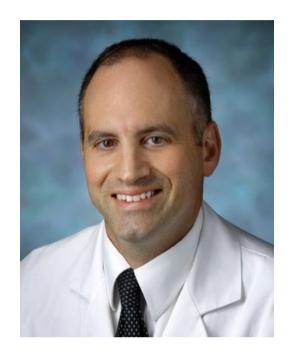
- physical activity and adiposity
- and hepatic triglyceride secretion



- Endocrine mediators secreted by adipose tissue - Regulation of fat mass, systemic insulin sensitivity, glucose, and lipid

- CTRP13 ablation improves systemic glucose and lipid metabolism - Hypermetabolism in mice carrying a near-complete human chromosome 21 - PRADC1: a novel metabolic-responsive secretory protein that modulates - <u>C1q/TNF-related protein 2 (CTRP2) deletion promotes adipose tissue lipolysis</u>

DIABETES, LIPID DISORDERS, ENDOCRINOLOGY TODD TARQUIN BROWN, MD, PHD



Faculty Webpage **Publications Grant Information** **APPOINTMENTS**

Professor, Medicine Member, Center for Metabolism and Obesity research

RESEARCH INTERESTS

- patients
- antiretroviral treatment

RESEARCH **HIGHLIGHTS**

- **Physical Function**
- Lipid Disorders in People with HIV





- Exploring metabolic, endocrine and skeletal abnormalities in HIV-infected

- Understanding the occurrence and prevalence of insulin resistance, diabetes, and anthropometric changes in HIV patients and their relationship to

- Skeletal muscle DNA methylation: Effects of exercise and HIV - CROI 2023: Metabolic and Other Complications of HIV Infection - Associations of Muscle Density and Area With Coronary Artery Plaque and

- The association of adipose tissue area with subclinical coronary atherosclerosis progression in men with and without HIV

LIPID METABOLISM, CHOLESTEROL, CANCER PETER ESPENSHADE, PHD



Faculty Webpage **Publications Grant Information** **APPOINTMENTS**

Professor, Cell Biology Associate Dean, Graduate Biomedical Education Member, Center for Metabolism and Obesity Research

RESEARCH INTERESTS

in mammalian cells by regulating cholesterol and fatty acid synthesis

- Cellular regulation of cholesterol homeostasis and adaptation to hypoxia - Sterol Regulatory Element Binding Protein (SREBP) control of lipid homeostasis

RESEARCH **HIGHLIGHTS**

- Targeting Stearoyl-CoA Desaturase in Solid Tumors
- Expanding roles for SREBP in metabolism
- the Endoplasmic Reticulum





- <u>Serum lipoprotein-derived fatty acids regulate hypoxia-inducible factor</u>

- Dipyridamole Inhibits Lipogenic Gene Expression by Retaining SCAP-SREBP in

ENDOCRINOLOGY, DIABETES, HYPERLIPIDEMIA, GONADAL DYSFUNCTION ADRIAN SANDRA DOBS, MD, MHS



APPOINTMENTS

Professor, Medicine

RESEARCH INTERESTS

- Diabetes and polycystic ovarian syndrome

Faculty Webpage **Publications Grant Information**

RESEARCH **HIGHLIGHTS**

- **Care Setting**
- **Examination Survey**
- women with polycystic ovary syndrome



- Understanding the relationship between sex hormones and heart disease - Male gonadal functional and hormone replacement therapy

- An Individualized Approach to Managing Testosterone Therapy in the Primary

- Analysis of cardiovascular risk factors associated with serum testosterone levels according to the US 2011-2012 National Health and Nutrition

- Pathophysiology, risk factors, and screening methods for prediabetes in

CARDIOVASCULAR DISEASE, INSULIN RESISTANCE JEFFERY HUNTER YOUNG, MD, MHS



Faculty Webpage **Publications Grant Information** **APPOINTMENTS**

Associate Professor, Medicine

RESEARCH INTERESTS

RESEARCH **HIGHLIGHTS**

- Mitochondrial DNA copy number and diabetes: the Atherosclerosis Risk in **Communities (ARIC) study**
- The Sex and Race Specific Relationship between Anthropometry and Body Fat **Composition Determined from Computed Tomography: Evidence from the** Multi-Ethnic Study of Atherosclerosis





Member, Center for Metabolism and Obesity Research

- Genetic epidemiology and novel risk factors for cardiovascular disease - The roles of hypertension, diabetes, and obesity on cardiovascular disease

CARDIOLOGY AND OBESITY LILI AYALA BAROUCH, MD



APPOINTMENTS

Professor, Medicine Director, Sports Cardiology Program Member, Center for Metabolism and Obesity research

RESEARCH INTERESTS

- nutritional/metabolic cues

Faculty Webpage **Publications Grant Information**

RESEARCH **HIGHLIGHTS**

- model of obesity
- stiffness induced by chronic intermittent hypoxia
- **myocytes**





- Understanding molecular mechanisms to sense and respond to

- Exploring novel neuron-specific enzyme function in metabolic processes

- Beneficial cardiac effects of caloric restriction are lost with age in a murine

- Restoring leptin signaling reduces hyperlipidemia and improves vascular

- Dependence of β3-adrenergic signaling on the adipokine leptin in cardiac

ADDITIONAL FACULTY WITH CARDIO-METABOLIC DISEASE AND OBESITY FOCUS



ENDOCRINOLOGY AND OBESITY **REXFORD S. AHIMA, MD, PHD**



Faculty Webpage Publications Research Grants

APPOINTMENTS

Professor, Medicine Director, Division of Endocrinology, Diabetes, and Metabolism, Johns Hopkins **University School of Medicine**

RESEARCH INTERESTS

- Pathogenesis of obesity and diabetes
- Relationship between energy stores and regulation of energy balance in brain - Understanding the role of adipokines, cytokines and myokines in targeting the brain and peripheral organs to control feeding and metabolism

RESEARCH **HIGHLIGHTS**

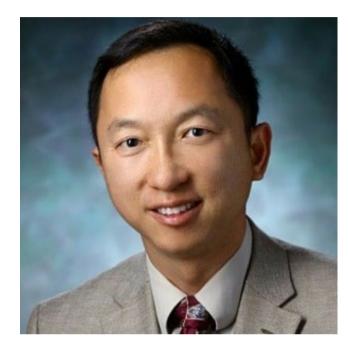
- resistance in primary mouse hepatocytes
- Endocrine disorders associated with obesity
- Feeding by CCR2 and CX3CR1.





- Inositol polyphosphate multikinase modulates free fatty acids-induced insulin - Synergistic Modulation of Inflammatory but not Metabolic Effects of High-Fat

METABOLIC DISORDERS, DIABETES DAX FU, PHD



Faculty Webpage **Publications Grant Information**

APPOINTMENTS

RESEARCH INTERESTS

Associate Professor, Physiology

- Structure and function of a human pancreatic zinc transporter for the early diagnosis and treatment of diabetes
- Role of zinc transporter ZnT8 in regulating insulin processing and secretion

RESEARCH **HIGHLIGHTS**

- Novel autoantibodies to the β-cell surface epitopes of ZnT8 in patients progressing to type-1 diabetes
- Down-regulation of the islet-specific zinc transporter-8 (ZnT8) protects human insulinoma cells against inflammatory stress
- Highly specific monoclonal antibodies for allosteric inhibition and immunodetection of the human pancreatic zinc transporter ZnT8
- <u>Coupling of Insulin Secretion and Display of a Granule-resident Zinc</u> Transporter ZnT8 on the Surface of Pancreatic Beta Cells.
- for Type-2 Diabetes.



- Lipid-tuned Zinc Transport Activity of Human ZnT8 Protein Correlates with Risk

METABOLISM, DIABETES LING HE, MD, PHD



Faculty Webpage **Publications Grant Information**

APPOINTMENTS

Associate Professor, Pediatrics

RESEARCH INTERESTS

- Liver metabolism of glucose and lipids
- Mitochondrial dynamics and respiration through AMPK activation
- Insulin resistance and hepatic steatosis in obesity and type II diabetes

RESEARCH **HIGHLIGHTS**

- Blocking AMPKaS496 phosphorylation improves mitochondrial dynamics and hyperglycemia in aging and obesity
- Mitochondrial Dynamics during Development
- the Insulin Receptor
- Activation of the canonical ER stress IRE1-XBP1 pathway by insulin regulates glucose and lipid metabolism
- The P300 acetyltransferase inhibitor C646 promotes membrane translocation of insulin receptor protein substrate and interaction with the insulin receptor
- <u>Alterations of Gut Microbiota by Overnutrition Impact Gluconeogenic Gene</u> **Expression and Insulin Signaling**



- Far-western Blotting Detection of the Binding of Insulin Receptor Substrate to

CARDIOVASCULAR DISEASE DAVID A. KASS, MD



Faculty Webpage Publications Grant Information

APPOINTMENTS

Professor, Medicine Abraham and Virginia Weiss Professor of Cardiology **Professor of Biomedical Engineering**

RESEARCH INTERESTS

- and heart failure
- Models of obesity, metabolic syndrome, and cardiac stress to identify treatments of heart failure with preserved ejection fraction
- Role of mToR signaling in cardiac disease, TRPC6 and muscular dystrophy
- Sarcomere enhancing molecules and pulmonary hypertension
- Transient receptor potential canonical type 6 (TRPC6) O-GlcNAcylation at Threonine-221 plays potent role in channel regulation
- Myocardial Metabolomics of Human Heart Failure With Preserved Ejection Fraction
- Pharmacological TRPC6 inhibition improves survival and muscle function in mice with Duchenne muscular dystrophy
- Single serine on TSC2 exerts biased control over mTORC1 activation mediated by ERK1/2 but not Akt
- Increased Energy Expenditure and Protection From Diet-Induced Obesity in Mice Lacking the cGMP-Specific Phosphodiesterase PDE9

RESEARCH **HIGHLIGHTS**



- Professor of Pharmacology and Molecular Sciences
- Protein kinase G in heart disease and treatment, Phosphodiesterase inhibitors

ENDOCRINOLOGY AND OBESITY ANASTASIA KRALLI, PHD



<u>Faculty Webpage</u> <u>Publications</u> <u>Research Grants</u> **APPOINTMENTS**

Professor, Physiology

RESEARCH INTERESTS

- Role of estrogen-related receptors in adipose tissue and skeletal muscle
- Therapeutic interventions for states where oxidative metabolism or tissue function are compromised
- Regulatory and transc adaptations

RESEARCH HIGHLIGHTS

- Loss of skeletal musc intolerance
- <u>Perm1 regulates CaMKII activation and shapes skeletal muscle responses to</u> <u>endurance exercise training</u>
- <u>Estrogen-Related Receptors Mediate the Adaptive Response of Brown Adipose</u> <u>Tissue to Adrenergic Stimulation.</u>
- <u>Complementary Roles of Estrogen-Related Receptors in Brown Adipocyte</u> <u>Thermogenic Function.</u>
- <u>Perm1 enhances mitochondrial biogenesis, oxidative capacity, and fatigue</u> <u>resistance in adult skeletal muscle.</u>



- Regulatory and transcriptional mechanisms that enable physiologic

- Loss of skeletal muscle estrogen-related receptors leads to severe exercise

ADVANCED HEART FAILURE, CARDIAC TRANSPLANT, CARDIOVASCULAR DISEASE PRIYA UMAPATHI, MD



Faculty Webpage **Publications**

APPOINTMENTS

Assistant Professor, Medicine

RESEARCH INTERESTS

- Cardiac metabolism, glycobiology and proteins - Therapeutic targets and precision medicine for heart failure - Inherited cardiomyopathies, cardio-metabolic heart failure - Cardiac remodeling/energetics and SGLT2 inhibitors - The Umapathi Cardio-Metabolic Lab

RESEARCH **HIGHLIGHTS**

- <u>CaMKII as a Therapeutic Target in Cardiovascular Disease</u> - Excessive O-GlcNAcylation Causes Heart Failure and Sudden Death - Differential Detection of O-GlcNAcylated proteins in the heart using antibodies



CARDIOVASCULAR DISEASE, LIPID DISORDERS, OBESITY ERIN MICHOS, MD, MHS

RESEARCH INTERESTS

APPOINTMENTS



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RESEARCH **HIGHLIGHTS** Associate Professor, Medicine Director, Women's Cardiovascular Health Research

- - Advanced Lipid Disorders

 - Obesity and Cardiometabolic Diseases

 - cardiovascular disease prevention in women
 - Atherosclerotic Cardiovascular Disease
 - of their kidney and heart protection
 - **Obesity**





- Cardio-Obstetrics and Cardiovascular Disease in Women

- Risk prediction for cardiovascular disease including the use of coronary artery calcium scores, markers of inflammation, and other biomarkers

- Cardiovascular disease prevention in women - the current state in 2023 - <u>Sex-specific differences in cardiovascular risk factors and implications for</u> - Inclisiran: A New Strategy for LDL-C Lowering and Prevention of - Glucagon-like peptide-1 receptor agonists in diabetic kidney disease: A review - Role of Glucagon-Like Peptide-1 Receptor Agonists in Achieving Weight Loss

and Improving Cardiovascular Outcomes in People With Overweight and

CARDIOVASCULAR DISEASE, RISK STRATIFICATION, DYSLIPIDEMIA, MI CHIADI ERICSON NDUMELE, MD, PHD, MHS



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APPOINTMENTS

RESEARCH INTERESTS

RESEARCH **HIGHLIGHTS** Associate Professor, Medicine Robert E. Meyerhoff Assistant Professor

- Cardio-metabolic disease
- Biomarkers and cardiovascular risk assessment
- **American Heart Association**

- Heart Failure and Obesity: The Latest Pandemic



- Relationship between obesity and cardiovascular disease

- For obese people, an increased risk of 'silent' heart damage, study says - Cardiovascular-Kidney-Metabolic Health: A Presidential Advisory From the

- Addressing Cardiovascular Risk in Diabetes: It's More Than the Sugar

- Obesity and Subtypes of Incident Cardiovascular Disease

Cardiovascular epidemiology; Vascular & Kidney diseases Josef Coresh, MD, PhD



Faculty Webpage
Publications
<u>Research Grants</u>

APPOINTMENTS	Professor, Epidemiology George W. Comstock Professor
RESEARCH INTERESTS	- Chronic Kidney Disease and the - Chronic Kidney Disease Prognos
RESEARCH HIGHLIGHTS	 Novel Prediction Equations for A Cardiovascular Disease Incorpo Health: A Scientific Statement Fr Cardiovascular-Kidney-Metaboli American Heart Association Polygenic Risk Scores for Kidney Circulating Proteome, and Incide Chronic Kidney Disease Testing A Diabetes Across 24 U.S. Health (



e Risk of Cardiovascular Disease osis Consortium

Absolute Risk Assessment of Total porating Cardiovascular-Kidney-Metabolic From the American Heart Association olic Health: A Presidential Advisory From the

ey Function and Their Associations with dent Kidney Diseases g Among Primary Care Patients With Type 2 h Care Organizations

TARGETED DRUG DELIVERY PLATFORMS KANNAN RANGARAMANUJAM, PHD, MS



Faculty Webpage **Publications Grant Information** **APPOINTMENTS**

RESEARCH

HIGHLIGHTS

RESEARCH INTERESTS

Professor, Ophthalmology Co-Director, Center for Nanomedicine

- Targeted drug delivery for ocular and neurodegenerative diseases
- Targeted drug delivery using dendrimers
- Galactosylated hydroxyl-polyamidoamine dendrimer targets hepatocytes and improves therapeutic outcomes in a severe model of acetaminophen poisoning-induced liver failure
- Rationally Designed Galactose Dendrimer for Hepatocyte-Specific Targeting and Intracellular Drug Delivery for the Treatment of Liver Disorders
- <u>Systemic administration of dendrimer N-acetyl cysteine improves outcomes</u> and survival following cardiac arrest



OTHER RELATED INSTITUTES AT JOHNS HOPKINS

- Johns Hopkins Heart and Vascular Institute
- Johns Hopkins Comprehensive Diabetes Center
- Endocrinology, Diabetes, and Metabolism Research



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