

Cardiovascular Research Institute

- Summer 2023 Newsletter -



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MARK L. ENTMAN, MD

On June 15th, we had the pleasure of gathering with colleagues and friends in the lobby of the DeBakey Museum at Baylor College of Medicine to celebrate Dr. Mark L. Entman's retirement. After 53 years of service at Baylor College of Medicine, Dr. Entman retired at the end of June 2023. Among his countless accomplishments, Dr. Entman's commitment to educational programs is unmatched. As we celebrate our esteemed colleague, we would like to highlight his background as well as memories over the years.

Dr. Entman was recruited to Baylor as an assistant professor in 1970. He was a Howard Hughes Medical Investigator from 1971-1979. In 1977, Dr. Entman became the Chief of the Section of Cardiovascular Sciences and the Director of the Division of Research of the NHLBI National Research and Demonstration Center (now the DeBakey Heart Center) at Baylor College of Medicine and The Methodist Hospital from 1976-1985. Dr. Entman has been an inspirational leader



whose research has spanned a range of topics, including the role of myocardial calcium and sarcoplasmic reticulum function, acute inflammation and myocardial injury, and the chronic inflammatory response in cardiac repair and remodeling. He received the Outstanding Research Award from the International Society of Heart Research (1986) and was awarded an NIH MERIT AWARD (1989-1999) for his work on sarcoplasmic reticulum calcium flux. Dr. Entman and his colleagues were the first to demonstrate the induction of ICAM-1 expression on viable cardiac myocytes in the jeopardized border zone of a reperfused myocardial infarction. They demonstrated in vivo that the presence of ICAM-1 sensitizes myocardial cells to adhesion-dependent neutrophil induced oxidative injury. Subsequent work dealt with characterizing the cytokine cascade responsible for inflammatory injury of the surviving border zone. Factors influencing both the cardiac and leukocyte response were studied and examined in the context of the pathophysiological course of infarct progress.

Before joining Baylor faculty, Dr. Entman's training at Duke University involved matriculation in the highly innovative Research Training Program designed to promote the proper background for cellular and molecular research for MD's seeking a career in academic medicine. In 1974, his former mentor at Duke, Dr. Salih Wakil, joined the Baylor faculty as chairman of biochemistry and the two collaborated in writing the NIH training grant to establish the MD/PhD Program at Baylor, of which Dr. Entman was a co-director until 1980. In 1978, Dr. Entman became the director of the Section of Cardiovascular Sciences in the Department of Medicine, and he was paramount in the new development of that program. The core curriculum for the DeBakey Heart Center Graduate Program arose from those efforts and was funded for many years by an NIH training grant which supported an independent graduate program directed by his colleague and close friend, Dr. Julius Allen. The



resources of this program also provided the structure of a Basic Science Training program in Pediatric Cardiology at Texas Children's Hospital which was financed by an independent NIH training program.

Dr. Entman has given countless lectures to trainees on the Cardiovascular Sciences PhD track and has been dedicated to furthering the educational mission at Baylor College of Medicine. He has mentored over 50 physician-scientists and researchers, many of whom are now leading cardiology departments and research programs across the US and world. His enthusiasm and commitment to the educational programs at Baylor College of Medicine is revered among his trainees and peers.

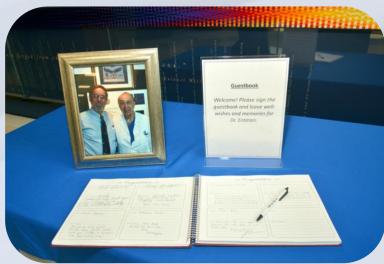


Pictured L to R: Drs. Xander Wehrens, Mark Entman, and Christie Ballantyne





Pictured L to R: Drs. William Zoghbi, Michael Schneider, Biykem Bozkurt, Mark Entman, A.J. Marian, and Christie Ballantyne





Pictured L to R: Sharon Malinowski, Dr. Mark Entman and Karima Ghazzaly



MEET PHYSICIAN-SCIENTIST AND CHIEF OF CARDIOLOGY AT BAYLOR COLLEGE OF MEDICINE, DR. REZA ARDEHALI

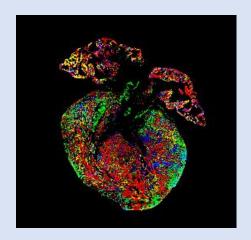
Reza Ardehali, MD, PhD was appointed Chief of Cardiology at Baylor College of Medicine in January 2023. He has had an impressive academic journey, driven by a passion for cardiology and a desire to advance the field through research, patient care, and program development.



Dr. Ardehali began his academic pursuits as an undergraduate at the University of Utah, where he majored in mathematics and chemistry. During this time, his interest in biomedical engineering led him to continue his education at the University of Utah, where he obtained a PhD degree in biomedical engineering with a focus on mechanical circulatory support, including left ventricular assist devices (LVADs) and total artificial heart. It was during this period that he developed a strong interest in cardiology and its potential for medical advancements. Following the completion of his PhD, Dr. Ardehali completed medical school at Emory University. Guided by his passion for cardiology, he pursued an internal medicine residency at The Johns Hopkins Hospital. To further specialize in the field, he then went on to complete a fellowship in cardiovascular medicine at Stanford University Medical Center. Driven by a fascination with heart regeneration, he joined the lab of Dr. Irving Weissman at Stanford, where he focused on the use of embryonic cells to understand heart

development and explore their potential for regenerative therapies.

Dr. Ardehali leads a dedicated team of researchers in his lab, which focuses on two main areas of research: using stem cells to explore cardiovascular development/regeneration and understanding the mechanisms that regulate cardiac fibrosis. His primary goal is to develop regenerative therapies that allow the heart to repair itself. This involves limiting fibrosis and promoting the regeneration of new cardiomyocytes, the muscle cells of the heart. The Ardehali lab is exploring the use of exogenous or endogenous cells to achieve this purpose. Additionally, they are investigating the mechanisms that regulate scar formation in the heart. They aim to identify the cell types responsible for fibrosis and understand the underlying signaling pathways involved. By gaining a deeper understanding of these processes, Dr. Ardehali hopes to develop diagnostic and treatment approaches to minimize scar formation in the heart.



Fluorescent microscope image of Mesp1^{Cre}; R26^{VT2/GK} (Rainbow mouse) heart at E14.5

As a physician-scientist, Dr. Ardehali is committed to providing high-quality patient care while pursuing cuttingedge research. He finds great satisfaction in mentoring colleagues and building successful programs. The opportunity to serve as the Section Chief of Cardiology at Baylor College of Medicine, an institution with a rich legacy in cardiovascular science and medicine, aligns perfectly with his goals of advancing patient care, conducting research, and engaging in administrative work to foster program development and mentorship. In his new role, Dr. Ardehali aims to enhance the organization of the cardiology section by fostering cohesive teams and subsections to improve the volume and value of clinical care provided.

Dr. Ardehali strongly believes in the power of collaboration and team science. He is actively seeking collaborations with other scientists and physician-scientists at Baylor College of Medicine, within the Texas Medical Center, and on an international and national scale. His goal is to integrate his work with others to achieve greater research outcomes. He brings a wealth of knowledge, experience, and a passion for cardiology to his role as Chief of Cardiology at Baylor College of Medicine. With a focus on advancing research, improving patient care, and building collaborative programs, he aims to make significant contributions to the field of cardiology while supporting the growth and success of his colleagues. His commitment to excellence and dedication to mentorship make him an invaluable asset to the BCM community.

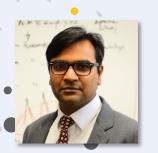
HONORS AND AWARDS



Dr. <u>David Durgan</u>, Assistant Professor of Integrative Physiology, has received the 2023 <u>Transformational Project Award</u> from the <u>American Heart Association</u>. The three-year, \$300,000 award supports his lab's research on the role of the gut microbiota on blood pressure regulation.







Dr. <u>Ketan Ghaghada</u>, Associate Professor of Radiology, has received the Innovator Award Program from <u>The Marfan Foundation</u>. The two-year, \$100,000 award supports his research on "Predicting of Aortic Dissection and Rupture in Marfan Syndrome Using Nano-Radiomics."

Additionally, he received the Innovative Project Award from American Heart Association. The two-year, \$200,00 award supports his research on the" Rational Design of Lipid Nanoparticles for Delivery of RNA Therapeutics to SMCs in Aortic Aneurysm and Dissection."



Dr. <u>Christie Ballantyne</u>, Chief, Cardiovascular Research, is ranked #560 in the world and #335 in the United States as a Top Scientist according to <u>Research.com</u>.





FACULTY AWARDS

Norton Rose Fulbright Faculty Award:



Dr. Waleed Kayani Assistant Professor, Cardiology



<u>Dr. Vijay Nambi</u> Associate Professor, Cardiology

Star Faculty Award for Excellence in Patient Care:



Dr. Ayse Mindikoglu
Associate Professor,
Gastroenterology and
Hepatology



Dr. Sai Kaumudi Saridey
Associate Professor,
Nephrology

Early Career Faculty Award for Excellence in Patient Care:



Dr. Pascale Khairallah
Assistant Professor,
Nephrology



<u>Dr. Bryan Tucker</u> Assistant Professor, Pulmonary, Critical Care, & Sleep

SEMINAR SERIES

The James T. Willerson, MD Cardiovascular Sciences Seminar Series continues in Fall 2023. In collaboration with Texas Heart Institute, these seminars are in-person and held at Baylor College of Medicine on select Wednesdays at noon. Visit the CVRI website for further details. CME credit is available to eligible attendees.



Sept. 6th

Katherine King, M.D., Ph.D.
Associate Professor
Department of Pediatrics-Infectious
Disease

Baylor College of Medicine

Seminar Location: McMillian Auditorium



Sept. 20th

Camilla Wenceslau, Ph.D., M.S., FAHA

Associate Professor

Cardiovascular Translational Research

Center

Department of Cell Biology and Anatomy School of Medicine Columbia

University of South Carolina

Seminar Location: McMillian Auditorium



Oct. 4th

Vijay Nambi, M.D.

Associate Professor Department of Medicine-Athero & Lipo

Baylor College of Medicine
Seminar Location: McMillian

Auditorium



Oct. 18th

Xue Gao, Ph.D.

T.N. Law Assistant Professor, Chemical and Biomolecular Engineering

Rice University

Seminar Location: 187A



Nov. 1st - Selected by trainee

Joseph Hill, M.D., Ph.D.

Professor of Medicine and Molecular Biology

James T. Willerson M.D. Distinguished Chair in Cardiovascular Diseases

Frank M. Ryburn, Jr. Chair in Heart Research

1100001011

UT Southwestern Medical Center **Seminar Location**: 187A



Nov. 8th – Mark L. Entman, MD Distinguished Lecture in Cardiovascular Research

Christopher Semsarian, MBBS, Ph.D.

Professor of Medicine, Sydney Medical School

Cardiologist, Royal Price Alfred Hospital, Central Clinical School Head, Molecular Cardiology Program, Centenary Institute

Seminar Location: Kleberg Auditorium



Nov. 29th

Yong Xu, M.D., Ph.D.

Professor, CNRC

Associate Director for Basic Sciences Department of Pediatrics and Nutrition

Baylor College of Medicine **Seminar Location**: Kleberg

Auditorium



Dec. 13th

Mingxia Gu, M.D., Ph.D., FAHA

Assistant Professor

Center for Stem Cell & Organoid

Medicine (CuSTOM)

Department of Pediatrics

Cincinnati Children's Hospital Medical

Center

Seminar Location: Kleberg Auditorium



American Heart Association Funding Opportunities

Opportunity	Proposal Deadline	Award Start Date
AHA Predoctoral Fellowship Enhances the training of promising students in pre-doctoral or clinical health professional degree training programs and who intend careers as scientists, physician-scientists or other clinician-scientists, or related	Sept. 6, 2023	Jan. 1, 2024
careers aimed at improving global health and wellbeing. AHA Postdoctoral Fellowship Enhances training of postdoctoral applicants who are not yet independent. The applicant must be embedded in an appropriate investigative group with the mentorship, support, and relevant scientific guidance of a research mentor.	Sept. 7, 2023	Jan.1, 2024
Career Development Award Supports highly promising healthcare and academic professionals in the early years of first professional appointment to assure the applicant's future success as a research scientist in the field of cardiovascular and/or cerebrovascular disease research.	Dec. 6, 2023	Apr. 1, 2024
Collaborative Sciences Award Fosters innovative collaborative approaches to research projects that propose novel pairings of investigators from at least two broadly disparate disciplines. The proposal must focus on the collaborative relationship, such that scientific objectives could not be achieved without the efforts of at least two co-principal investigators and their respective disciplines.	ТВА	TBA
Established Investigator Award Supports established investigators in rapid career growth phase, with records of accomplishments and showing extraordinary	ТВА	TBA
Innovative Project Award Supports highly innovative, high-impact research that could ultimately lead to critical discoveries or major advancements that will accelerate the field of cardiovascular and/or cerebrovascular research.	TBA	TBA
Research Supplement to Promote Diversity in Science Under the mentorship of current AHA awardees, this mechanism supports research experiences for predoctoral and postdoctoral fellows from underrepresented racial and ethnic groups in science.	TBA	TBA

See more funding opportunities through the American Heart Association, $\underline{\mathsf{HERE}}.$



Additional Funding Opportunities

Thoracic Surgery Foundation	Application Open	Application Deadline
TSF Research Award→ Operational support of original research efforts by cardiothoracic surgeons who have completed their formal training, and who are seeking initial support and recognition for the research program. Awards of up to \$85,000 per year for up to two years are granted to support the work of an early-career cardiothoracic surgeon (within seven years of first faculty appointment at time of application deadline).	Jul. 1, 2023	Sept. 15, 2023
STS Research Award → Operational support of original research efforts by cardiothoracic surgeons who have completed their formal training, and who are seeking initial support and recognition for the research program. Awards of up to \$90,000 per year for up to two years are granted to support the work of an early-career cardiothoracic surgeon (within seven years of first faculty appointment at time of application deadline). <i>The STS Research Award designation</i> is given to the highest-ranking TSF Research Award application.	Jul. 1, 2023	Sept. 15. 2023
TSF Resident Research Fellowship Award → This award provides up to \$60,000 per year for up to two years to support the research fellowship of a resident who has not yet completed cardiothoracic surgical training. During the fellowship, the resident will work in a cardiothoracic surgical clinical or laboratory research program.	Jul. 1, 2023	Sept. 15, 2023
Nina Starr Braunwald, MD was the first woman to be certified by the ABTS and the first woman to conduct open heart surgery. This award in her name provides operational support of original research efforts by women cardiac surgeons who have completed their formal training, and who are seeking initial support and recognition for their research program. Since its inception, award recipients have gone on to become established leaders within the field and this award is one of the specialty's most prestigious research grants. Awards of up to \$85,000 per year for up to two years are made each year to support the work of an early-career woman cardiac surgeon (within five years of first faculty appointment).	Jul. 1, 2023	Sept. 15, 2023
Nina Starr Braunwald Research Fellowship Award→ Nina Starr Braunwald, MD was the first woman to be certified by the ABTS and the first woman to conduct open heart surgery. This award in her name supports up to \$60,000 per year for up to two years for a woman resident working in a cardiac surgical clinic or laboratory research program who has not yet completed cardiothoracic surgical training.	Jul. 1, 2023	Sept. 15, 2023



Grants & Funding	Deadline
PAR-21-038	Sept. 26, 2023
Stephen I. Katz Early Stage Investigator Research Project Grant (R01	
Clinical Trial Not Allowed)	
Global Cardiovascular Research Funders Forum	Sept. 29, 2023
	, ,
PAR-18-771 I PAR-18-772	Oct. 12, 2023
NHLBI Career Transition Award for Intramural Postdoctoral Fellows and	
Research Trainees (K22)	
PA-20-187 I PA-20-188	Oct. 12, 2023
NIH Pathway to Indepence Award (Parent K99/R00)	, i
DAD 22 404	0 17 2022
PAR-22-194 NHLBI TOPMed: Omics Phenotypes of Heart, Lung, and Blood DIsorders	Oct. 17, 2023
(X01)	
(102)	
RFA-HL-24-005	Oct. 13, 2023 (New); Nov. 13,
Limited Competition: Small Grant Program for NHLBI K01/K08/K23/K25	2023
Recipients (R03 Clinical Trial Optional)	(Renewal/Resubmission/Revision)
Standard NIH Due Dates	

EDUCATION

Graduate School of Biomedical Science Cardiovascular Courses

The Cardiovascular Research Institute offers 3 graduate school courses open to PhD students, postdocs, trainees, staff, and all others interested in cardiovascular science. In the AY 23-24 academic year, the CVRI courses have been moved up a term.

Lectures are in person. Mid-term and Final Exams are provided via BlackBoard (for those taking the course for credit). For those interested in these courses, there are 3 ways to sign up and participate.

- Enroll with the GSBS for full credit and grade (tests, surveys and attendance required).
- Audit the course through GSBS for credit (surveys and attendance required for credit; no tests or letter grade assigned).
- View the CVRI recorded lectures of interest within the course date (survey participation required).
 Email <u>cvri@bcm.edu</u> for further details.

CVRI is committed to supporting the growth and success of our next generation of innovative scientists.

Term 2: GS-DD-6210 - Cardiovascular Diseases GS-DD-6210

2 Credits

Registration: September 11, 2023 - September 22, 2023

This course provides a general overview of the main, common cardiovascular diseases and their causes. Topics covered include atherosclerosis, hypertension, congenital heart disease, ischemic heart disease, cerebral stroke, cardiac arrhythmias, and the effects of aging on the cardiovascular system. Tuesday, Thursday | October 9, 2023 – December 8, 2023

Course Director: Xander Wehrens, MD, PhD

Term 3: GS-DD-6403 - Advanced Topics in Cardiac Pathophysiology and Disease

3 Credits

Registration: November 20, 2023 - December 1, 2023

This course covers the fundamentals of cardiac development, and cardiac function in both physiological and pathological conditions. The course will also discuss the cutting-edge research approaches used in cardiovascular research. <u>Tuesday</u>, <u>Wednesday</u>, <u>Thursday</u> | <u>January 2</u>, <u>2024 – March 1</u>, <u>2024</u>

Course Directors: Na Li, PhD & Xander Wehrens, MD, PhD

Term 4: GS-DD-6404 - Advanced Topics in Vascular Pathophysiology and Disease

3 Credits

Registration: February 12, 2024 - February 23, 2024

This course emphasizes cardiovascular disease pathology with a focus on vascular disorders and atherosclerosis. Lectures will cover all components of the normal system, inherited forms of disease, and the pathogenesis of acquired types of disease. Topics include vascular diseases, lipid disorders, atherosclerosis, hemostasis and bleeding disorders, microcirculation disorders, stroke, hypertension, and peripheral artery disease. The course will also discuss the cutting-edge research approaches used in cardiovascular research. Tuesday, Wednesday, Thursday | March 11, 2024 – May 10, 2024

Course Directors: William Lagor, PhD & Xander Wehrens, MD, PhD

CARDIOVASCULAR RESEARCH INSTITUTE 2023-2024 Pilot Awards

The Cardiovascular Research Institute (CVRI) at Baylor College of Medicine is pleased to announce pilot awards for proposals in cardiovascular research that will promote collaboration between investigators and are likely to lead to externally funded grants in the foreseeable future.

Timeline:

- Submit your proposal <u>HERE</u>.
 Deadline for submission is August 25, 2023.
- Funding for selected proposals is anticipated to be available October 2023- June 30, 2024.

Goal:

To promote the writing of multi-PI grants in the area of cardiovascular research (clinical, translational or basic). The expectation is that before the end of calendar year 2023, the co-PIs will submit a grant proposal for external funding (NIH preferred).

Eligibility:

- BCM faculty who are members of the CVRI may submit applications. Please visit: https://www.bcm.edu/research/research-centers/cardiovascular-research-institute/membership to become a member.
- Each proposal must include, but not exceed two PIs, with primary appointments in at least two different BCM departments. Collaborators from other TMC institutions are allowed.
- IRB or IACUC approval must be approved at the time of application submission
- Only BCM faculty and their lab may receive CVRI pilot funds.

Application:

- Specific aims page
- Max of 6 pages of research plan with: significance, innovation, experimental design, timeline
- Biosketches of the PIs and other essential personnel
- Budget details
- Specific proposal for obtaining extramural funding

Amount of funding:

• 3-4 awards up to \$20,000 will be made in unrestricted funds.

SELECT PUBLICATIONS

Share your work! Increase your impact!

Email cvri@bcm.edu with your latest publications and we will share it with our CV community on social media.

April 2023

Seyerle AA, Laurie CJ, Coombes B, Jain DP, Conomos, M, Brody J, Chen M-H M, Gogarte SM, Beutel K, Gupta NR, Heckbert SD, Jackson RD, Johnson A, Ko DE, Manson J, McKnight BA, Metcalf GC, Morrison AP, Reiner A, Sofer T, Tang WL, wiggins K, Boerwinkle E, de Andrade MB, Gabriel SA, **Gibbs RC**, Laurie CM, Psaty BS, Vasan R, Rice K, Kooperberg CS, Pankow JL, Smith N, Pankratz N. Whole Genome Analysis of Venous Thromboembolism: the Trans-Omics for Precision Medicine Program. Circ Genom Precis Med; 16(2): e003532.

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Ballantyne CM, Varughese MG, **Abushamat LA**. Lipid-Lowering Therapy in the Elderly: Are Current Guidelines a Sign of Ageism in Medical Care? J Am Coll Cardiol; 81(14): 1350-1352.

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May 2023

M. Directo cardiac reprogramming: A new technology for cardiac repair. J Mol Cell Cardiol; 178: 51-58.

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Khalid U, Kayani W, Alam M, Denktas AE. Revascularization Options for Left Main Disease: What Clinicians Need to Know. Curr Atherscler Rep; 25(6): 267-273.

Ouzounian M, **LeMaire SA**, **Milewicz DM**. Heritable aortic root aneurysms. Ann Cardiothorac Surg; 12(3): 265.267.

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Krittanaw ong C, Rodriguez M, Lui M, **Misra A**, Tang WHW, **Bozkurt B**, Yancy CW. Misconceptions and Facts about Heart Failure with Reduced Ejection Fraction. Am J Med; 136(5): 422-431.

June 2023

El-Ayash H, Puyau M, **Bacha F**. Hyperglycemia: A determinant of cardiac autonomic dysfunction in youth with obesity across the spectrum of glycemic regulation. Pediatr Obes; Online ahead of print.

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Zea-Vera R, Ryan CT, Navarro SM, Havelka J, Wall MJ, **Coselli JS**, **Rosengart TK**, **Chatterjee S**, **Ghanta RK**. Development of a Machine Learning Model to Predict Outcomes and Cost After Cardiac Surgery. Ann Thorac Surg; 115(6): 1533-1542.

Ayyaswamy S, Shi H, Zhang B, Bryan RM Jr, **Durgan DJ.** Obstructive Sleep Apnea-Induced Hypertension is Associated with Increased Gut and Neuroinflammation. JAHA; 12(11): e029218.

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Wang X, Song J, Yuan Y, Li L, Abu-Taha I, Heijman J, Sun L, Dobrev S, Kamler M, Xie L, Wehrens XHT, Horrigan FT, Dobrev D, Li N. Downregulation of FKBP5 Promotes Atrial Arrhythmogenesis. Circ Res; 133(1): e1-e16.

Kim JA, Kim SE, Ellenbogen KA, Vijayaraman P, **Chelu MG**. Clinical outcomes of conduction system pacing versus biventricular pacing for cardiac resychronization therapy: A systematic review and meta-analysis. J Cardiovasc Electrophysiol; Online ahead of print.

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July 2023

Segar MW, Zhang A, Paisley RD, Badjatiya A, Lambeth KD, Mullins K, **Razavi M**, Molina-Razavi JE, Rasekh A, Saeed M. Risk Stratification in Patients Who Underwent Percutaenous Left Atrial Appendage Occlusion. AM J Cardiol; Online ahead of print.

Krittanawong C, Qadeer YK, Hayes RB, Wang Z, **Virani SS**, Zeller M, Dadvand P, Lavie CJ. Noise Exposure and Cardiovascular Health. Curr Probl Cardiol; Online ahead of print.

Brown KN, Phan H, Jui EL, Kang MK, Connell JP, **Keswani SG**, **Grande-Allen KJ**. Isolation and Characterization of Porcine Endocardial Endothelial Cells. Tissue Eng Part C Methods; Online ahead of print.

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EXECUTIVE LEADERSHIP COMMITTEE



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Biykem Bozkurt, MD, PhD CVRI Associate Director



Reza Ardehali, MD, PhD Medicine, Cardiology



Christie Ballantyne, MD Medicine, Atherosclerosis & Lipoprotien



Changyi Johnny Chen MD, PhD Surgery, Vascular Surgery



Mihail G. Chelu, MD, PhD Medicine, Cardiology



Katarzyna Cieslik, PhD Medicine, Cardiovascular Science



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James Martin, MD, PhD Integrative Physiology



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