

Heart Disease Fact Sheet

CIRM funds many projects seeking to better understand heart disease and to translate those discoveries into new therapies.

Description

Heart disease strikes in many forms, but collectively it causes one third of all deaths in the U.S. Many forms of heart disease have a common result—cardiomyopathy. While this is commonly called congestive heart failure (CHF), it is really just the heart becoming less efficient due to any number of causes, but the most common is loss of functioning heart muscle due to the damage caused by a heart attack. An estimated 4.8 million Americans have CHF, with 400,000 new cases diagnosed each year. Half die within five years.

Numerous clinical trials are underway testing a type of stem cell found in bone marrow, called mesenchymal stem cells or MSCs, to see if they are effective in treating the form of CHF that follows a heart attack. While those trials have shown some small improvements in patients the researchers have not found that the MSCs are creating replacement heart muscle. They think the improvements may be due to the MSCs creating new blood vessels that then help make the existing heart muscle healthier, or in other ways strengthening the existing tissue.

California's stem cell agency has numerous awards looking into heart disease (the full list is below). Most of these involve looking for ways to create stem cells that can replace the damaged heart muscle, restoring the heart's ability to efficiently pump blood around the body. Some researchers are looking to go beyond transplanting cells into the heart and are instead exploring the use of tissue engineering technologies, such as building artificial scaffolds in the lab and loading them with stem cells that, when placed in the heart, may stimulate the recovery of the muscle.

Other CIRM-funded researchers are working in the laboratory, looking at stem cells from heart disease patients to better understand the disease and even using those models to discover and test new drugs to see if they are effective in treating heart disease. Other researchers are trying to make a type of specialized heart cell called a pacemaker cell, which helps keep a proper rhythm to the heart's beat.

We also fund projects that are trying to take promising therapies out of the laboratory and closer to being tested in people. In some cases, these awards also fund the early phase clinical trials to show that they are safe to use and, in some cases, show some signs of being effective.

Clinical Stage Programs

Cedars-Sinai Medical Center

Pulmonary arterial hypertension (PAH) is a progressive condition with no cure. Scientists at Cedars-Sinai Medical Center are using donor cells derived from the heart to reduce two hallmark symptoms of pulmonary hypertension: inflammation and high blood pressure in the blood vessels within the lungs. These conditions make the heart struggle to pump blood to the heart and lungs, and over time, can ultimately lead to heart failure. The aim of this treatment is to delay the progression of the disease.

- To learn more about this clinical trial
- To read about this trial on clinicaltrials.gov

Capricor (Heart failure and Duchenne Muscular Dystrophy-related heart failure)

Capricor is using donor cells derived from heart stem cells developed by Cedars-Sinai to treat patients developing heart failure after a heart attack. In early studies the cells appear to reduce scar tissue, promote blood vessel growth and improve heart function.

Read about the team's progress

In a second trial, Capricor is using the same donor cells derived from heart stem cells to treat patients developing heart failure due to

Duchenne Muscular Dystrophy. In early studies the cells appear to reduce scar tissue, promote blood vessel growth and improve heart function.

Read about the team's progress

Cedars-Sinai Medical Center (Cardiomyopathy)

This team developed a way to isolate some heart-specific stem cells that are found in adult heart muscle. They use clumps of cells called Cardiospheres to reduce scarring caused by heart attacks. Initially they used cells obtained from the patient's own heart but they later developed methods to obtain the cells they need from donor organs, which allows the procedure to become an off-the-shelf-therapy, meaning it can be available when and where the patient needs it rather than having to create it new each time. The company, working with the Cedars-Sinai team, received FDA approval to begin a clinical trial in June 2012.

• Read about the team's progress

Stanford School of Medicine (Heart Failure)

This team plans to turn embryonic stem cells into what are called cardiomyocytes, the kind of cells that can become heart muscle. They plan to develop methods for producing sufficient quantities for clinical therapy and to do all the laboratory work and preliminary testing needed to gain FDA approval of a clinical trial by the close of the grant. They are proposing to carry out a trial with patients who have disease that is so advanced that they are on a waiting list for heart transplants.

•	Read	about	the	team's	progress

Video: Bruce Conklin of the Gladstone Institute of Cardiovascular Disease talks about using stem cells to screen drugs for heart side effects

CIRM Grants Targeting Heart Disease

Researcher name	Institution	Grant Title	Grant Type	Award Amount
Christian Metallo	University of California, San Diego	Metabolic regulation of cardiac differentiation and maturation	Basic Biology V	\$1,124,834
Brian Black	University of California, San Francisco	Weinstein Cardiovascular Development Conference	Conference	\$35,000

PILAR RUIZ- LOZANO	Regencor, Inc.	IND-enabling Studies of Wearable Evolve-FSTL1 for Cardiac Regeneration after MI	Therapeutic Translational Research Projects	\$3,923,191
Irving Weissman	Stanford University	Prospective isolation of hESC-derived hematopoietic and cardiomyocyte stem cells	Comprehensive Grant	\$2,471,386
Deepak Srivastava	Gladstone Institutes, J. David	Direct Cardiac Reprogramming for Heart Regeneration	Early Translational III	\$5,795,871
Eduardo Marbán	Cedars-Sinai Medical Center	[REDACTED] Heart Disease Regenerative Medicine Team Planning Award	Disease Team Planning	\$38,980
Alice Tarantal	University of California, Davis	11th Annual Gene Therapy Symposium for Heart, Lung, and Blood Diseases	Conference	\$16,850
Randall Lee	University of California, San Francisco	Embryonic Stem Cell-Derived Therapies Targeting Cardiac Ischemic Disease	Comprehensive Grant	\$2,424,353
Joseph Wu	Stanford University	Heart Repair with Human Tissue Engineered Myocardium	Early Translational III	\$4,396,738
Mark Mercola	Sanford Burnham Prebys Medical Discovery Institute	"Stem Cell Therapies for Heart Failure"	Disease Team Planning	\$44,450
Alice Tarantal	University of California, Davis	gth Annual Gene Therapy Symposium for Heart, Lung and Blood Diseases	Conference	\$12,000
Harold Bernstein	University of California, San Francisco	Modeling Myocardial Therapy with Human Embryonic Stem Cells	Comprehensive Grant	\$2,134,694
Walter Boyd	University of California, Davis	Extracellular Matrix Bioscaffold Augmented with Human Stem Cells for Cardiovascular Repair	Early Translational III	\$4,631,754
Michelle Khine	University of California, Merced	Micro Platform for Controlled Cardiac Myocyte Differentiation	SEED Grant	\$193,700
Alice Tarantal	University of California, Davis	10th Annual Gene Therapy Symposium for Heart, Lung, and Blood Diseases	Conference	\$18,300
Robb Maclellan	University of California, Los Angeles	Human Cardiovascular Progenitors, their Niches and Control of Self-renewal and Cell Fate	Basic Biology I	\$917.667
Christopher Zarins	Stanford University	Engineering a Cardiovascular Tissue Graft from Human Embryonic Stem Cells	Comprehensive Grant	\$2,454,490
Eric Adler	University of California, San Diego	Identification of Novel Therapeutics for Danon Disease Using an iPS Model of the Disease	Early Translational III	\$1,701,575

Todd McDevitt	Gladstone Institutes, J. David	Engineering microscale tissue constructs from human pluripotent stem cells	Research Leadership	\$5,849,100
Eduardo Marbán	Cedars-Sinai Medical Center	Autologous cardiac-derived cells for advanced ischemic cardiomyopathy	Disease Team Research I	\$5,560,232
Deepak Srivastava	Gladstone Institutes, J. David	microRNA Regulation of Cardiomyocyte Differentiation from Human Embryonic Stem Cells	Comprehensive Grant	\$2,994,719
Yang Xu	University of California, San Diego	Human ES cell based therapy of heart failure without allogenic immune rejection	Early Translational III	\$1,857,600
Rachel Smith	Capricor, Inc	Allogeneic Cardiac-Derived Stem Cells for Patients Following a Myocardial Infarction	Disease Team Therapy Development - Research	\$14,405,857
Joseph Gold	City of Hope, Beckman Research Institute	Development of a scalable, practical, and transferable GMP-compliant suspension culture-based differentiation process for cardiomyocyte production from human embryonic stem cells.	Tools and Technologies III	\$891,518
Sonja Schrepfer	University of California, San Francisco	Hypo-immunogenic cardiac patches for myocardial regeneration	Inception - Discovery Stage Research Projects	\$235,818
Bruce Conklin	Gladstone Institutes, J. David	Induced Pluripotent Stem Cells for Cardiovascular Diagnostics	New Cell Lines	\$1,708,560
Mark Mercola	Sanford Burnham Prebys Medical Discovery Institute	Chemical Genetic Approach to Production of hESC-derived Cardiomyocytes	Comprehensive Grant	\$2,832,000
Joseph Wu	Stanford University	Human Embryonic Stem Cell-Derived Cardiomyocytes for Patients with End Stage Heart Failure	Disease Team Therapy Development - Research	\$19,060,330
Shaochen Chen	University of California, San Diego	Development of 3D Bioprinting Techniques using Human Embryonic Stem Cells Derived Cardiomyocytes for Cardiac Tissue Engineering	Tools and Technologies III	\$1,368,264
Kevin Healy	University of California, Berkeley	Human Cardiac Chip for Assessment of Proarrhythmic Risk	Quest - Discovery Stage Research Projects	\$899,581
Michael Longaker	Stanford University	Derivation and analysis of pluripotent stem cell lines with inherited TGF-b mediated disorders from donated IVF embryos and reprogrammed adult skin fibroblasts	New Cell Lines	\$1,406,636
Patrick McDonough	Vala Sciences, Inc.	Optimization in the Identification, Selection and Induction of Maturation of Subtypes of Cardiomyocytes derived from Human Embryonic Stem Cells	Tools and Technologies I	\$870,717

Huei-sheng Chen	Sanford Burnham Prebys Medical Discovery Institute	Studying Arrhythmogenic Right Ventricular Dysplasia with patient-specific iPS cells	Basic Biology IV	\$1,579,250
Joseph Wu	Stanford University	Macaca mulatta as advanced model for predictive preclinical testing of engineered cardiac autografts and allografts	Tools and Technologies III	\$1,689,744
Mark Mercola	Stanford University	Multipotent Cardiovascular Progenitor Regeneration of the Myocardium after MI	Quest - Discovery Stage Research Projects	\$1,809,234
Sylvia Evans	University of California, San Diego	Specification of Ventricular Myocyte and Pacemaker Lineages Utilizing Human Embryonic Stem Cells	SEED Grant	\$585,600
Mohammad Pashmforoush	University of Southern California	Transcriptional Regulation of Cardiac Pacemaker Cell Progenitors	New Faculty I	\$2,816,578
Benoit Bruneau	Gladstone Institutes, J. David	Epigenetic regulation of human cardiac differentiation	Basic Biology IV	\$1,568,148
Joseph Wu	Stanford University	Drug Discovery & Stem Cell Models for Cardiovascular Disease Conference	Conference II	\$7.500
Deborah Lieu	University of California, Davis	Microenvironment for hiPSC-derived pacemaking cardiomyocytes	Quest - Discovery Stage Research Projects	\$2,042,438
Krishna Shenoy	Stanford University	Technology for hESC-Derived Cardiomyocyte Differentiation and Optimization of Graft-Host Integration in Adult Myocardium	SEED Grant	\$572,891
Andrew Putnam	University of California, Irvine	A Novel Engineered Niche to Explore the Vasculogenic Potential of Embryonic Stem Cells	New Faculty I	\$395,764
Deborah Lieu	University of California, Davis	Induction of Pluripotent Stem Cell-Derived Pacemaking Cells	Basic Biology IV	\$1,333,689
Linda Cambier	Cedars-Sinai Medical Center	Exosomal Y-RNAs as mediators of bioactivity of cardiacderived cell therapy	Inception - Discovery Stage Research Projects	\$181,063
Jane Lebkowski	Geron Corporation	Preclinical Development and First-In-Human Testing of GRNCM1 in Advanced Heart Failure	Disease Team Therapy Planning	\$o
John Cashman	Human BioMolecular Research Institute	Discovering Potent Molecules with Human ESCs to Treat Heart Disease	SEED Grant	\$688,274
Irving	Stanford University	Antibody tools to deplete or isolate teratogenic, cardiac, and blood stem cells from hESCs	Tools and Technologies II	\$1,463,814

Sheng Ding	Gladstone Institutes, J. David	A new paradigm of lineage-specific reprogramming	Basic Biology IV	\$1,568,395
Phillip Yang	Stanford University	Activation of patient-specific endogenous myocardial repair through the exosomes generated from the hypoxic iPSC-derived cardiomyocytes (iCMs).	Inception - Discovery Stage Research Projects	\$234,619
Eric Adler	University of California, San Diego	Genetically Modified Hematopoietic Stem Cells for the Treatment of Danon Disease	Quest - Discovery Stage Research Projects	\$1,393,200
Huei-sheng Chen	Sanford Burnham Prebys Medical Discovery Institute	Development of Neuro-Coupled Human Embryonic Stem Cell-Derived Cardiac Pacemaker Cells.	SEED Grant	\$695,680
Deepak Srivastava	Gladstone Institutes, J. David	Mechanisms of Direct Cardiac Reprogramming	Basic Biology III	\$1,572,380
Eduardo Marbán	Cedars-Sinai Medical Center	Mechanism of heart regeneration by cardiosphere-derived cells	Basic Biology IV	\$1,367,604
Arjun Deb	University of California, Los Angeles	Targeting progenitors in scar tissue to reduce chronic scar burden	Inception - Discovery Stage Research Projects	\$230,400
Joseph Wu	Stanford University	A Phase I, Pilot Study of Human Embryonic Stem Cell- Derived Cardiomyocytes in PaTients with ChrOnic Ischemic Left VentRicular Dysfunction (HECTOR)	Clinical Trial Stage Projects	\$6,987.507
Michelle Khine	University of California, Irvine	Micro Platform for Controlled Cardiac Myocyte Differentiation	SEED Grant	\$156,426
Ali Nsair	University of California, Los Angeles	Characterization and Engineering of the Cardiac Stem Cell Niche	Basic Biology III	\$1,127,741
Ali Nsair	University of California, Los Angeles	Human Induced Pluripotent Stem Cell-Derived Cardiovascular Progenitor Cells for Cardiac Cell Therapy.	New Faculty Physician Scientist	\$2,316,894
Deepak Srivastava	Gladstone Institutes, J. David	Use of Human iPSC-derived Endothelial Cells for Calcific Aortic Valve Disease Therapeutics	Quest - Discovery Stage Research Projects	\$2,400,048
Deborah Lieu	University of California, Davis	Building a hiPSC-based biopacemaker	Quest - Discovery Stage Research Projects	\$1,260,827
Phillip Yang	Stanford University	In Vivo Molecular Magnetic Resonance Imaging of Human Embryonic Stem Cells in Murine Model of Myocardial Infarction	SEED Grant	\$629,952

Diego Junction Maturation and Disease Using Human IPSC Reza Ardehalti University of California, Los Angeles Preclinical evaluation of human embryonic stem cellderived cardiovascular progenitors in a large animal model Scientist Stage Projects Angeles University Physician Scientist Stage Projects Stanford University Division of Cardiosphere-Derived Cells for Duchenne Muscular Dystrophy Cardiomyopathy Benoit Gladstone Induction of Cardiosphere-Derived Cells for Duchenne Muscular Dystrophy Cardiomyopathy Benoit Gladstone Induction of Cardiospensis in pluripotent cells via chromatin remodeling factors Benoit Institutes J David Induction of Cardiogenesis in pluripotent cells via chromatin remodeling factors Benoit Institutes J David Induction of Cardiogenesis in pluripotent cells via chromatin remodeling factors Benoit Institutes J David Induction of Cardiogenesis in pluripotent cells via chromatin remodeling factors Benoit Institutes J David Induction of Cardiogenesis in pluripotent cells via chromatin Rew Faculty II S2,723.6 Benoit Institutes J David Induction of Cardiogenesis in pluripotent cells via chromatin Rew Faculty II S2,723.6 Benoit Institutes J David Induction of Cardiogenesis in pluripotent cells via chromatin Rew Faculty II S2,723.6 Benoit Institute Cardiomyopathy with Human Induced Pluripotent Stem Cells Institute					
Reza Ardehali California. Los Angeles derived cardiovascular progenitors in a large animal model Physician Scientist	Farah Sheikh	California, San	, -	Basic Biology III	\$1,341,955
Linda Marban Capricor, Inc Muscular Dystrophy Cardiomyopathy Stage Projects S3,376.2 Phillip Yang Stanford University Hypoxia-specific Production of Exosomes from IPSC- derivatives for Myocardial Repair Induction of cardiogenesis in pluripotent cells via chromatin Induction Induction of cardiogenesis in pluripotent cells via chromatin Induction of Disease Research in Cardiovascular Diseases Induction of Disease Research in Cardiovascular Diseases Induction of Induction of Induction of Induction of VEGF Producing MSC for the Treatment of Critical Limb Ischemia Induction of Cardiovascular Projects Induction of VEGF Producing MSC for the Treatment of Critical Limb Ischemia Induction of Regenerative Medicine Induction of Projects Ind	Reza Ardehali	California, Los	,	Physician	\$2,930,388
Phillip Yang Stanford University derivatives for Myocardial Repair Stage Research Projects Stage Research Projects Induction of Cardiogenesis in pluripotent cells via chromatin New Faculty II \$2.723.6 Induction of Cardiogenesis in pluripotent cells via chromatin New Faculty II \$2.723.6 Sean Wu Stanford University Elucidating Molecular Basis of Hypertrophic Cardiomyopathy with Human Induced Pluripotent Stem Cells Stanford University Tissue Collection for Accelerating IPSC Research in Cardiovascular Diseases Modeling Molecular Basis of Hypertrophic Cardiovascular Diseases Modeling S1.291.8; Michael Lewis Cedars-Sinai Medical Center Cardiovascular Diseases Modeling Stanford University A novel hybrid CRISPR tool for gene network perturbation And hipSC engineering PsC Research Projects S704.66 Projects Stanford University Physiology and disease New Faculty II S3.005.6 University Physiology and disease New Faculty II S4.005.6 Institute Improving Existing Drugs for Long QT Syndrome type 3 (LOT3) by hipSC Disease-in-Dish Model Projects S70.361.3 Improving Existing Drugs for Long QT Syndrome type 3 (LOT3) by hipSC Disease-in-Dish Model Projects S70.361.3 Institute Institutes J. David Direct Cardiac Reprogramming for Regenerative Medicine Institutes J. David Targeting stromal progenitors to prevent the development Therapeutic Translational Research Projects Annelse Stanford Projects S4.841.4 Research Annelse S4.841.4 Research S4.841.4 Res	Linda Marban	Capricor, Inc	,		\$3,376,259
Bruneau Institutes, J. David remodeling factors Sean Wu Stanford University Cardiomyopathy with Human Induced Pluripotent Stem Cells Stanford University Stanford Cells Stanford University Tissue Collection for Accelerating iPSC Research in Cardiovascular Diseases Modeling Michael Lewis Medical Center Pulmonary Arterial Hypertension Treated with Cardiovascular Diseases Modeling Stanford University A novel hybrid CRISPR tool for gene network perturbation and hiPSC engineering Ching-Pin Chang University VEGF signaling in adventitial stem cells in vascular physiology and disease John Laird University of California, Davis Treatment of Critical Limb Ischemia Deepak Srivastava Gladstone Institutes, J. David University of Cardiox Reprogramming for Regenerative Medicine Search Projects Targeting stromal progenitors to prevent the development of heart failure Stanted University of Cardiornia, Los Angeles Angeles Angeles Angeles Angeles Angeles Angeles Sangeles	Phillip Yang			Stage Research	\$1,418,023
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Stanley Qi Stanford University A novel hybrid CRISPR tool for gene network perturbation and hiPSC engineering Stanford University VEGF signaling in adventitial stem cells in vascular physiology and disease University of California, Davis Human BioMolecular Cashman Gladstone Institutes, J. David Arjun Deb Stanford University A novel hybrid CRISPR tool for gene network perturbation and hiPSC engineering VEGF signaling in adventitial stem cells in vascular physiology and disease New Faculty II \$3,005,6 Phase I study of IM Injection of VEGF Producing MSC for the Treatment of Critical Limb Ischemia Improving Existing Drugs for Long QT Syndrome type 3 (LQT3) by hiPSC Disease-in-Dish Model Institutes Direct Cardiac Reprogramming for Regenerative Medicine California, Los Angueles Targeting stromal progenitors to prevent the development of heart failure Stage Projects \$7,34.7 Quest - Discovery Stage Research Projects \$4,841.4	Joseph Wu			for Disease	\$1,291,832
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BioMolecular Research Improving Existing Drugs for Long QT Syndrome type 3 Early Translational IV \$6,361,31	John Laird	,			\$76,066
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Arjun Deb California, Los Angeles Targeting stromal progenitors to prevent the development of heart failure Targeting stromal progenitors to prevent the development of heart failure \$4,841,41	•		Direct Cardiac Reprogramming for Regenerative Medicine	Stage Research	\$2,392,157
	Arjun Deb	California, Los		Translational Research	\$4,841,428
Kara McCloskey University of California, Merced Building Cardiac Tissue from Stem Cells and Natural Matrices New Faculty II \$1,656,0		California,		New Faculty II	\$1,656,083

Robert Robbins	Stanford University	Human Embryonic Stem Cell-Derived Cardiomyocytes for Patients with End Stage Heart Failure	Disease Team Therapy Planning I	\$73,030	
Joseph Wu	Stanford University	A Novel, Robust and Comprehensive Predictive Tool Using Human Disease-Specific Induced Pluripotent Stem Cells for Preclinical Drug Screening	Tool Translational Research Projects	\$975,000	
					Total: \$172,735,333

CIRM Heart Disease Videos



Growing Stem Cell Research in California: Todd McDevitt Lab, Gladstone Institutes



Heart disease: Progress toward stem cell therapies, a live Google Hangout



Webinar: Stem Cell Clinical Trial for Heart Disease - Lessons Learned | Eduardo Marbán



Deepak Srivastava, Gladstone Institutes - CIRM Stem Cell #SciencePitch



Spotlight on Stem Cell Advances in Pediatric Heart Disease: A Change of Heart



Spotlight on Stem Cell Advances in Heart Disease: Generating Muscle within an Existing Heart



Joshua's Heart Story: Pediatric Heart Disease and the Promise of Stem Cell Advances



Stem Cell Clinical Trial for Heart Failure: Eduardo Marbán -CIRM Spotlight on Disease



Cardiovascular Therapies: Spotlight on Stem Cell Research - Welcoming Remarks



Cardiovascular Therapies: Spotlight on Stem Cell Research -W. Douglas Boyd



Cardiovascular Therapies: Spotlight on Stem Cell Research - Yung-Wei Chi



Cardiovascular Therapies: Spotlight on Stem Cell Research - Jeanette Owens



Valentine's Day Stem Cell Wish: Mending Broken Hearts



Stem Cell-Derived Heart Cells: Bruce Conklin - CIRM Science Writer's Seminar

Resources

- Blogs on heart disease research from the CIRM Stem Cellar
- NIH: Heart Failure Information
- Find a clinical trial near you: NIH Clinical Trials database
- American Heart Association
- · National Heart, Lung and Blood Institute

• CDC: Heart Disease

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