# Jonathan Rodolfo Silva, PhD FAHA

## **Professor**

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Department of Biomedical Engineering

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## **Positions**

Department of Biomedical Engineering

Washington University in St. Louis

Professor 2023 - present
Dennis & Barbara Kessler Career Development Associate Professor 2021 - 2023
Associate Professor 2018 - 2023
Assistant Professor 2012 - 2018

Department Director of Diversity 2020 – present

Department of Computer Science and Engineering, Adjunct 2020 - present

SentiAR, Inc.

Founder, Director and Chief Technology Officer 2017 – present

Excera, Inc.

Founder, Director and Chief Technology Officer 2021 – present

## **Education**

Washington University in St. Louis, St. Louis, MO

PhD in Biomedical Engineering. 2008

Case Western Reserve University, Cleveland, OH

MS in Biomedical Engineering. 2004

The Johns Hopkins University, Baltimore, MD

BS in Biomedical Engineering. 2000

## **Pre- and Postdoctoral Research**

#### Washington University in St. Louis, St. Louis, MO

Postdoctoral training. Department of Cell Biology. Colin Nichols, PhD. 2011 to 2012

Measured and modeled physiological effects of K,ATP heterogeneity in the heart and the pancreas.

#### University of Chicago, Chicago, IL

Postdoctoral training. Department of Pediatrics. Steven AN Goldstein MD PhD. 2008 to 2011 Probed Na<sup>+</sup> channel slow inactivation using simultaneous cut-open oocyte recording and fluorescence.

## Washington University in St. Louis, MO

Doctoral program, Department of Biomedical Engineering. Yoram Rudy, PhD 2004 to 2008 Dissertation: Linking Ion-Channel Molecular Dynamics to the Action Potential: Cardiac I<sub>Ks</sub> Simulations.

#### Case Western Reserve University, Cleveland, OH

MS program, Department of Biomedical Engineering. Yoram Rudy, PhD 2000 to 2004

Thesis: Subunit interaction determines Iks Participation in Cardiac Repolarization and Repolarization Reserve.

# **Honors and Awards**

- 2022. American Heart Association Established Investigator Award
- 2018. Fellow American Heart Association
- 2011. Burroughs Wellcome Fund Career Award at the Scientific Interface
- 2007. AAAS/Science Sponsored Membership
- 2005. Cardiac Arrhythmias Gordon Conference: Best Poster
- 2003. Housing and Residence Life Distinguished Service Award, Case Western Reserve University
- 2002. Graduate Dean's Award for Instructional Excellence, Case Western Reserve University
- 1999. Biomedical Engineering Honor Society (Alpha Eta Mu Beta), The Johns Hopkins University
- 1996. National Merit Scholar

#### Honors to students:

- 2019. Kathryn Mangold, Best Graduate Poster, Cardiovascular Research Day
- 2019. Kuo Chan Weng, Best Postdoctoral Poster, Cardiovascular Research Day
- 2018. Emily Wagner, Best Poster, CIMED Research Day
- 2018. Wandi Zhu, Washington University Biomedical Engineering Research Award
- 2018. Wandi Zhu, Philip Needleman Pharmacology Prize
- 2018. Wandi Zhu, Biophysical Society Student Research Achievement Award
- 2017. Wandi Zhu, Best Presentation, Gordon Research Seminar
- 2016. Caroline Dong, LMI Aerospace Award for Excellence in Research
- 2015. Wandi Zhu, Best Poster, Cardiovascular Research Day
- 2015. Wandi Zhu, Best Translational Poster, Bo Koster Memorial Symposium
- 2015. Bianca Yu, LMI Aerospace Award for Excellence in Research
- 2014. Eric Hsu, Barry M. Goldwater Scholar
- 2014. Ruben Garcia, Best Pre-College Poster AGMUS Research Symposium

# **Fellowships**

- 2011. Burroughs Wellcome Fund, Career Award at the Scientific Interface
- 2011. NIH Cardiovascular Biology Training Grant Washington University School of Medicine
- 2008. NIH Cardiovascular Pathophysiology and Biochemistry Training Grant University of Chicago
- 2001. NIH NRSA Pre-doctoral Fellowship
- 2000. Case Prime Fellowship
  - Case Western Reserve University
- 1999. Provost's Award for Undergraduate Research The Johns Hopkins University
- 1998. Research Experience for Undergraduates (REU) Fellowship
  - University of California, San Diego/San Diego Supercomputing Center
- 1997. Ford Foundation Award for Minority Research
  The Johns Hopkins University

# **Research Funding**

Current as PI or MPI

American Heart Association Established Investigator Award \$400,000 7/1/22-6/30/27

Precision Antiarrhythmic Medicine: Using Biophysical Insight to Improve Therapy

Developing molecules that target the cardiac Na<sup>+</sup> channel voltage sensing domains to improve efficacy of class I antiarrhythmics and assess the role of amiodarone metabolites in determining efficacy.

NIH-NHLBI, R01HL150637

\$2,292,835

7/1/20-6/30/24

Post-Transcriptional Regulation of Myocardial Sodium Channels

Collaborative project to understand how molecular level regulation of Nav channels by intracellular fibroblast growth factors determines myocyte electrophysiology.

NIH-NHLBI, R01HL148803

\$3,169,663

4/1/20-3/31/25

Personalizing Class I anti-Arrhythmic Drug Therapy

Goal is to leverage biophysical insight to predict whether patients with ventricular tachycardia or fibrillation will respond to class I anti-arrhythmic molecules.

NIH-NHLBI, R01HL142301

\$2,149,137

4/1/19-3/31/23 (NCE)

Mechanism of BK Channel Gating

Goal is to delineate mechanisms of the dual regulation of BK channels by membrane potential and intracellular Ca<sup>2+</sup>.

NIH-NHLBI, 1R01HL136553

\$1,906,250

4/1/17-3/31/23 (NCE)

Modulation of Cardiac NaV Channel Function and Electrophysiology by Interacting Molecules

Combines advanced molecular imaging modalities with multi-scale modeling to understand how molecules that interact with cardiac NaV channels affect arrhythmia susceptibility.

Current as Co-I or Preceptor

NIH-NIDDK, T32 DK108742

06/2016-05/2021

Imaging, Modeling and Engineering of Diabetic Tissues

Designed to prepare trainees with engineering backgrounds for research focused on fundamental mechanisms and translational possibilities for diabetes and metabolic diseases.

Role: Preceptor

NIH-NHLBI, T32 HL134635

7/1/17-6/30/22

Training in Integrative and Systems Biology of Cardiovascular Disease

Provide research opportunities; fundamental education and basic training in cardiovascular physiology.

Role: Preceptor

Previous as PI or MPI

NIH-NHLBI, 1R44HL140896

\$2,221,372

2/1/18-7/31/21

Using Augmented Reality to Make Cardiac Ablation Procedures Simpler and Safer

SBIR FastTrack grant is to SentiAR, Inc., a WashU spinout, whose goal is to commercialize an augmented reality platform that was developed in the Silva lab for use in the cardiac catheterization laboratory.

Institute of Clinical and Translational Sciences, 733

\$3,990

10/7/20-10/6/21

Post-translational Mechanisms in the Coordinated Regulation of Cardiac Rhythms

Create mouse models of FGF regulation of Nav channels

Children's Discovery Institute, CH- II-2017-575 \$450,000

Improving Outcomes In The Pediatric Cardiac Catheterization Lab Using Augmented Reality

Using mixed reality to guide cardiac ablation (year 1) and catheterization (year 2&3) procedures.

NIH-NHLBI, R01NS092570

\$333,594

3/1/20 - 2/28/21

2/1/17-1/31/21

KCNQ Channels: Gating and Subunits Modulation

Goal is to understand how intracellular and transmembrane subunits regulate KCNQ channel gating.

NIH, UG3 TR002170-01 7/1/17-4/30/20

A 3D In-Vitro Disease Model of Atrial Conduction

Aim is to create a compact system to test new therapies for atrial fibrillation using adenoviral delivery.

Role: Co-I

Skandalaris Center LEAP Inventor Challenge

\$50,000

1/1/19-12/31/19

Using Merged Reality to Improve Cochlear Implant Outcomes

Leveraging our experience in creating augmented reality solutions for cardiac ablation to improve cochlear implant outcomes.

CIMED Center Pilot Grant

\$25,000

1/1/18-12/31/18

Effects of Mechanical Forces on Cardiac Excitability in Heart Disease

Pilot grant in collaboration with Michael Greenberg and Jonathan Moreno developing technology to simultaneously assess excitability and contraction in iPSC-derived cardiomyocytes.

American Heart Association, 16GRNT31200025

\$154,000

7/1/16-6/30/18

Molecular Pathology of Brugada Syndrome Mutations

Goal is to use molecular imaging methods to discover the molecular basis for arrhythmia.

School of Engineering Collaboration Initiation Grant

\$25,000

4/25/17-3/24/18

Applying Mixed Reality to Otolaryngology Procedures

Using an augmented reality display to assist surgeons who are performing otolaryngology procedures.

Bear Cub Grant \$55,000 6/1/16-5/31/17

Improving Transcatheter Ablation Outcomes with Virtual Reality

Goal is to used mixed reality to enhance visualization during catheter ablation procedures.

Burroughs Wellcome Fund, 1010299

\$500,000

7/1/11-6/30/17

Applying molecular spectroscopy to derive multi-scale cardiac bioelectricity models

Goal is to obtain experimental data regarding the molecular motions of the cardiac Na+ channel.

Institute for Clinical and Translational Sciences, JIT410

\$5,000

7/11/16-7/10/17

The Nav Channel Beta Subunit Role in Chamber-Specific Heart Pharmacology

Editing iPSC-derived myocytes to test beta subunits alter Class 1b drug pharmacology.

CIMED Center Pilot Grant

\$25,000

1/1/14-12/31/14

Fluorescent Unnatural Amino Acid Tracking of Membrane Protein Conformation

Develop protocols to use fluorescent un-natural amino acids for voltage-clamp.

CIMED Center Pilot Grant

\$25,000

1/1/14-12/31/14

ATP binding to the cytosolic domain of KCNQ1 channel proteins

Goal is to assess mechanisms of ATP binding to KCNQ1 ion channels.

**Grants to Trainees** 

NIH F31 to Lucy Woodbury \$270,516 9/1/23-8/30/26

Investigating the interactions of auxillary subunits with the Nav1.5 channel

NIH F30 to Druv Bhagavan \$268,838 4/1/22-3/31/25

Connecting Subcellular Electrophysiology to Patient Arrhythmia: A Case Study of Amiodarone

American Heart Association Fellowship to Druv Bhagavan \$64,072 7/1/15-6/30/17

Connecting Molecular Electrophysiology to Patient Arrhythmia: A Case Study of Amiodarone

NIH K08 to Jonathan D. Moreno MD PhD \$745,988 7/1/21-6/30/26

3-Dimensional virtual ventricles to design precision therapies in hypertrophic cardiomyopathy

American Heart Association Fellowship to Wandi Zhu \$52,000 7/1/15-6/30/17

Discovering molecular mechanisms of how β-subunits regulate cardiac Na<sup>+</sup> Channels

CBSE Center Scholarship to Wandi Zhu \$10,000 7/1/15-6/30/16

American Heart Association Fellowship to Bicong Li \$4,000 7/1/16-9/1/16

HHMI Summer Research Fellowship (x5) \$15,000 Summer 2013-22

Student(s): Chen Zhao, Eric Hsu, Linxuan Yang, Namit Sambare, Sushruth Manchineella

American Heart Association Scholarship to Alex Goldberg \$2000 Summer 2016

## **Publications**

#### **Peer Reviewed Publications**

- Lesage A, Lorenzini M, Burel S, Sarlandie M, Bibault F, Lindskog C, Maloney D, *Silva JR*, Townsend RR, Nerbonne JM, Marionneau C. Determinants of iFGF13-mediated regulation of myocardial voltagegated sodium (NaV) channels in mouse. *Journal of General Physiology*. 2023 Sep 4;155(9). [PMID: 37516919]
- Bloom D, Catherall D, Miller N, Southworth MK, Glatz AC, Silva JR\*, Silva JN\*. Use of a Mixed Reality System for Navigational Mapping During Cardiac Electrophysiology Testing Does Not Prolong Case Duration: A Sub-analysis from the Cardiac Augmented Reality Study. Cardiovascular Digital Health Journal. 2023 Jun 14. [PMID: 37600447] \*co-corresponding author
- 3. Angsutararux P, Dutta AK, Marras M, Abella C, Mellor RL, Shi J, Nerbonne JM\*, *Silva JR*\*. Differential regulation of cardiac sodium channels by intracellular fibroblast growth factors. *Journal of General Physiology*. 2023 Mar 21;155(5):e202213300. [PMID: 36944081] \*co-corresponding author
- 4. Hanson A, McClenaghan C, Weng KC, Colijn S, Stratman AN, Halabi CM, Grange DK, *Silva JR*, Nichols CG. Electrophysiology of human iPSC-derived vascular smooth muscle cells and cell autonomous consequences of Cantu Syndrome mutations. *bioRxiv*. 2023:2023-06. [PMID: 37425756]
- 5. Moreno JD, *Silva JR*, Emerging methods to model cardiac ion channel and myocyte electrophysiology. *Biophys Rev (Melville)*. 2023 Mar;4(1):011315 [PMID: 37034130]
- 6. Hernandez B, Roberts B, Jamro-Comer E, Kodidhi A, Roelle L, Miller N, Littell L, Ybarra A, *Silva JR*, Silva JNA, Orr W. Validity Study Comparing Polar Ignite's® Estimated VO2max to Traditional

- Cardiopulmonary Exercise Testing in Normal Cardiac Anatomy & Fontan Pediatric Patients. *American Journal of Pediatrics*. 2023;9(2):79-84. [PMID: Pending]
- 7. Kang PW, Woodbury L, Angsutararux P, Sambare N, Shi J, Marras M, Abella C, Bedi A, Zinn D, Cui J, *Silva JR*. Arrhythmia-associated Calmodulin Variants Interact with KCNQ1 to Confer Aberrant Membrane Trafficking and Function. *bioRxiv*. 2023:2023-01.
- 8. Zhang G, Xu X, Jia Z, Geng Y, Liang H, Shi J, Marras M, Abella C, Magleby KL, *Silva JR\**, Chen J\*, Zou X\*, Cui J\*. An allosteric modulator activates BK channels by perturbing coupling between Ca2+ binding and pore opening. *Nature Communications*. 2022 Nov 9;13(1):6784. [PMID: 36351900] \*co-corresponding author
- 9. Bloom D, Colombo JN, Miller N, Southworth MK, Andrews C, Henry A, Orr WB, *Silva JR\**, Silva JN\*. Early preclinical experience of a mixed reality ultrasound system with active GUIDance for NEedlebased interventions: The GUIDE study. *Cardiovascular Digital Health Journal*. 2022 Oct 1;3(5):232-40. [PMID: 36310686] \*co-corresponding author
- 10. Oguntuyo K, Schuftan D, Guo J, Simmons D, Bhagavan D, Moreno JD, Kang PW, Miller E, *Silva JR*, Huebsch N. Robust, Automated Analysis of Electrophysiology in iPSC-Derived Micro-Heart Muscle for Drug Toxicity. *Tissue Engineering Part C*. 2022 Jun 7 (online). [PMID: 35925789]
- 11. Mangold KE, Zhou Z, Schoening M, Moreno JD, *Silva JR*. Creating Ion Channel Kinetic Models Using Cloud Computing. *Current Protocols*. 2022 Feb;2(2):e374. [PMID: 35175690]
- 12. Ransdell JL, Bhagavan D, *Silva JR*, Nerbonne JM. Intrinsic mechanisms in the gating of resurgent Na+currents. *eLife*. 2022 Jan 25;11:e70173. [PMID: 35076394]
- 13. Moreno JD, Bhagavan D, Li A, Gerstner NC, Miller EW, Huebsch N, Cresci S, *Silva JR*. Pulsus Alternans in Cardiogenic Shock Recapitulated in Single Cell Fluorescence Imaging of a Patient's Cardiomyocyte. *Circulation: Heart Failure*. 2021 Dec 10. [PMID: 34886677]
- 14. Angsutararux P, Zhu W, Voelker T, *Silva JR*. Molecular pathology of sodium channel beta-subunit variants. *Frontiers in Pharmacology*. 2021 Nov 19:3220. [PMID: 34867379]
- 15. Jung C, Wolff G, Wernly B, Bruno R, Franz M, Schulze PC, Silva JNA, *Silva JR*, Bhqatt D, Kelm M, "Virtual and augmented reality in cardiovascular care: State of the art and future perspectives", *JACC:* Cardiovascular Imaging, 2021 Mar;15(3):519-532. [PMID: 34656478]
- Mangold KE, Wang W, Johnson EK, Bhagavan D, Moreno JD, Nerbonne JM, Silva JR. "Identification of Structures for Ion Channel Kinetic Models". PLoS Computational Biology. 2021 Aug 17. [PMID: 34398881]
- 17. Zhu W, Wang W, Angsutararux P, Mellor RL, Isom LL, Nerbonne JM, *Silva JR*. "Modulation of the effects of Class-Ib antiarrhythmics on cardiac NaV1. 5-encoded channels by accessory NaVβ subunits", *Journal of Clinical Investigation: Insight*. 2021 Jun 22. [PMID: 34156986]
- 18. Gholami Derami H, Gupta P, Weng KC, Seth A, Gupta R, *Silva JR*, Raman B, Singamaneni S. "Reversible Photothermal Modulation of Electrical Activity of Excitable Cells using Polydopamine Nanoparticles". *Advanced Materials*. 2021 Jul 3:2008809. [PMID: 34216406]
- 19. Angsutararux P, Kang PW, Zhu W, *Silva JR*. "Conformations of voltage-sensing domain III differentially define NaV channel closed-and open-state inactivation." *Journal of General Physiology*. 2021 Aug 4;153(9):e202112891. [PMID: 34347027]
- Prakosa A, Southworth MK, Silva JN, Silva JR, Trayanova NA. "Impact of augmented-reality improvement in ablation catheter navigation as assessed by virtual-heart simulations of ventricular tachycardia ablation." Computers in Biology and Medicine. 2021 Jun 1; 133:104366. [PMID: 33836448]

- 21. Silva JN, Southworth MK, Andrews CM, Privitera MB, Henry AB, *Silva JR*. "Design Considerations for Interacting and Navigating with 2 Dimensional and 3 Dimensional Medical Images in Virtual, Augmented and Mixed Reality Medical Applications." In *International Conference on Human-Computer Interaction* 2021 Jul 24 (pp. 117-133). Springer, Cham. [PMID: 35079751]
- 22. Lorenzini M, Burel S, Lesage A, Wagner E, Charrière C, Chevillard M, Evrard B, Maloney D, Ruff K, Pappu R, Wagner S, Nerbonne J, *Silva JR*, Townsend RR, Maier LS, Marionneau C, "Proteomic and functional mapping of cardiac Nav1.5 channel phosphorylation", *Journal of General Physiology*, 2021 153 (2): e202012646. [PMID: 33410863]
- 23. Andrews CM, Henry AB, Soriano IM, Southworth MK, *Silva JR*, "Registration Techniques for Clinical applications of 3-Dimensional Augmented Reality Devices", *IEEE Journal of Translational Engineering in Health and Medicine*, 2020 Dec 17. [PMID: 33489483]
- 24. Kang PW, Westerlund AM, Shi J, White KM, Dou AK, Cui AH, *Silva JR*, Delemotte L, Cui J, "Calmodulin acts as a state-dependent switch to control a cardiac potassium channel opening", *Science Advances*, 2020 Dec 11; 6(50):eabd6798. [PMID: 33310856]
- 25. Southworth MK, Silva JNA, Blume W, Van Hare GF, Dalal AS, *Silva JR*, Performance Evaluation of Mixed Reality Display for Guidance during Transcatheter Cardiac Mapping and Ablation, *IEEE Journal of Translational Engineering in Health and Medicine*, 2020 Jul 3. [PMID: 32742821]
- 26. Zhu W, Li T, *Silva JR*, Chen J. Conservation and divergence in NaChBac and NaV1.7 pharmacology reveals novel drug interaction mechanisms. *Scientific Reports*. 2020 Jul 1;10(1):1-1. [PMID: 32612253]
- 27. Silva JN, Southworth MK, Blume WM, Andrews C, Van Hare GF, Dalal AS, Miller N, Sodhi SS, *Silva JR*. First-In-Human Use of a Mixed Reality Display During Cardiac Ablation Procedures. *Journal of the American College of Cardiology: Clinical Electrophysiology*. 2020 Jul 8. [PMID: 32819517]
- 28. Silva JN, Privitera MB, Southworth MK, *Silva JR*. Development and Human Factors Considerations for Extended Reality Applications in Medicine: The Enhanced Electrophysiology Visualization and Interaction System (ĒLVIS). *International Conference on Human-Computer Interaction* 2020 Jul 19 (pp. 341-356). Springer, Cham. [PMID: 34327520]
- 29. Salvage SC, Zhu W, Habib ZF, Hwang SS, Irons JR, Huang CL\*, *Silva JR\**, Jackson AP\*. Gating control of the cardiac sodium channel Nav1. 5 by its β3-subunit involves distinct roles for a transmembrane glutamic acid and the extracellular domain. *Journal of Biological Chemistry*. 2019 Dec 20;294(51):19752-63. \*Co-corresponding authors. [PMID: 31659116]
- 30. Moreno JD, Zhu W, Mangold K, Chung W, *Silva JR*. A molecularly detailed Nav1. 5 model reveals a new class I antiarrhythmic target. *Journal of the American College of Cardiology: Basic to Translational Science*. 2019 Oct 28;4(6):736-51. [PMID: 31709321]
- 31. Southworth, MK, *Silva JR\**, Silva JNA\*. Use of Extended Realities in Cardiology. *Trends in Cardiovascular Medicine*. Apr. 2020. \*Co-corresponding authors. [PMID: 31076168]
- 32. Andrews C, Southworth MK, Silva JNA\*, *Silva JR*\*. Extended Reality In Medical Practice. *Current Treatment Options in Cardiovascular Medicine*. 2019 Apr 1;21(4):18 . \*Co-corresponding authors [PMID: 30929093]
- 33. Zhu W, Mazzanti A, Voelker T, Hou P, Moreno JD, Angsutararux P, Naegle KM, Priori SG, *Silva JR*, Predicting Patient Response to the Antiarrhythmic Mexiletine Based on Genetic Variation: Personalized Medicine for Long QT Syndrome. *Circulation Research* 2019;124:539:552. [PMID: 30566038]
- 34. Silva JNA, Southworth M, Raptis C, *Silva JR*, "Emergence of Virtual and Augmented Reality in Cardiology", *Journal of the American College of Cardiology: Basic to Translational Science*, 3(3),420-430, 2018. [PMID: 30062228]

- 35. McJunkin JL, Jiramongkolchai P, Chung W, Southworth M, Durakovic N, Buchman CA, *Silva JR*. Development of a Mixed Reality Platform for Lateral Skull Base Anatomy. *Otology & Neurotology*. Dec 1;39(10):e1137-42, 2018. [PMID: 30239435]
- 36. *Silva JR* "How to Connect Cardiac Excitation to the Atomic Interactions of ion channels", *Biophysical Journal*, 114(2),259-266, 2018. [PMID: 29401425]
- 37. Peters CH, Yu A, Zhu W, *Silva JR*, Ruben PC, "Depolarization of the Conductance-Voltage Relationship in the Na<sub>V</sub>1.5 Mixed Syndrome Mutant, E1784K, Caused by Modified Fast Inactivation", *PLoS One*, 12(9):e0184605, 2017 [PMID: 28898267]
- 38. Zhu W, Voelker TL, Varga Z, Schubert AR, Nerbonne JM, *Silva JR* "Mechanisms of non-Covalent β Subunit Regulation of Nav Channel Gating", *J of General Physiology,* August; 149(8), 2017. [PMID: 28720590]
- 39. Mangold K, Brumback B, Angsutararux P, Zhu W, *Silva JR* "Mechanisms and Models of Cardiac Na<sup>+</sup> Channel Inactivation", *Channels*, 2017. [PMID: 28837385]
- 40. Hsu EJ, Zhu W, Schubert AR, Voelker T, Varga Z, *Silva JR* "Regulation of Na+ Channel Inactivation by the DIII and DIV Voltage-Sensing Domains", *J of General Physiology,* March; 149(3):389, 2017. [PMID: 28232510]
- 41. Teed ZR, *Silva JR* "A Computationally Efficient Algorithm for Fitting Ion Channel Gating Parameters", *MethodsX*, 3, 577-588, 2016. [PMID: 27924282]
- 42. Zhu W, Varga Z, Silva JR. Molecular motions that shape the cardiac action potential: Insights from voltage clamp fluorometry. *Progress in biophysics and molecular biology*. 2016 Jan 1;120(1-3):3-17. [PMID: 26724572]
- 43. Wang HG, Zhu W, Kanter RJ, *Silva JR*, Honeywell C, Gow RM, Pitt GS "A Novel NaV1.5 Voltage Sensor Mutation Associated with Severe Atrial and Ventricular Arrhythmias", *Journal of Molecular and Cellular Cardiology*, March; 92:52-62, 2016. [PMID: 26801742]
- 44. Brettmann J, Urusova D, Tonelli M, *Silva JR*, Henzler-Wildman K. "Role of Protein Dynamics in Ion Selectivity and Allosteric Coupling in the NaK Channel", *Proceedings of the National Academy of Science*, 112(50):15366-15371, 2015. [PMID: 26621745]
- 45. Varga, Z, Zhu W, Schubert AR, Pardieck JL, Hsu EJ, Zaydman MA, Cui J, *Silva JR*. "Direct Measurement of Cardiac Na<sup>+</sup> Channel Conformations Reveals Molecular Pathologies of Inherited Mutations." *Circulation: Arrhythmia and Electrophysiology*, 8:1228-1239, 2015. [PMID: 26283144]
- 46. Rudokas MW, Varga Z, Asaro AB, Schubert AR, *Silva JR*. "The *Xenopus* Oocyte Cut-Open Vaseline Gap Voltage-Clamp Technique with Fluorometry" *J of Visualized Exp*, e51040-e51040, 2014. [PMID: 24637712]
- 47. *Silva JR*, Cooper P, Nichols CG, "Modeling K,ATP-dependent Excitability in Pancreatic Islets" *Biophysical Journal*. 107(9), 2016-2026, 2014. [PMID: 25418087]
- 48. *Silva JR*. "Slow inactivation of Na<sup>+</sup> channels. Voltage Gated Sodium Channels". *Handbook of Experimental Pharmacology*. Feb. 2014 vol. 221, 33-49.
- 49. Zaydman MA, *Silva JR*, Delaloye K, Larsson HP, Cui J. "Kv7. 1 ion channels require a lipid to couple voltage sensing to pore opening" *Proceedings of the National Academy of Science*, August;110(32):13180-13185, 2013. [PMID: 23151230]
- 50. [Zhang HX\*, *Silva JR*\*], Lin YW, Verbsky JW, Lee US, Kanter EM, Yamada KA, Schuessler RB, Nichols CG. "The Heterogeneity and Function of KATP Channels in Canine Hearts" *Heart Rhythm*, \*Both authors contributed equally, October; 10(10):1576-1583, 2013. [PMID: 23871704]

- 51. *Silva JR* and Goldstein SAN. "Voltage Sensor Movements Describe Slow Inactivation of Na<sub>V</sub> Channels I: Wild Type" *Journal of General Physiology*. March; 141:309-321, 2013. [PMID: 23401571]
- 52. *Silva JR* and Goldstein SAN. "Voltage Sensor Movements Describe Slow Inactivation of Na<sub>V</sub> Channels II: L689I Mutants" *Journal of General Physiology*. March; 141:323-334, 2013. [PMID: 23401572]
- 53. Zaydman MA, *Silva JR*, Cui J. "Ion Channel Associated Diseases: Overview of Molecular Mechanisms" *Chemical Reviews*. November; 112(12)6319-6333, 2012. [PMID: 23151230]
- 54. Silva JNA and *Silva JR*. "Updates on the inherited cardiac ion channelopathies: from cell to clinical" *Current Treatment Options in Cardiovascular Medicine* October; 14(5):473-489, 2012. [PMID: 22865245]
- 55. *Silva JR* and Rudy Y. "Multi-scale Electrophysiology Modeling: From Atom to Organ" *Journal of General Physiology*. May; 135:575-581, 2010. [PMID: 20513759]
- 56. Silva JR, Pan H, Wu D, Nekouzadeh A, Decker K, Cui J, Baker N, Sept D, Rudy Y. "A Multiscale Model Linking Ion-Channel Molecular Dynamics and Electrostatics to the Cardiac Action Potential" Proceedings of the National Academy of Science. June; 106 (27):11102-11106, 2009. [PMID: 19549851]
- 57. Decker K, Heijman J, *Silva JR*, Hund T, Rudy Y. "Properties and Ionic Mechanisms of Action Potential Adaptation, Restitution and Accommodation in Canine Epicardium" *Am J Physiol: Heart* 2009 Jan; 296:1017-1026, 2009. [PMID: 19168720]
- 58. Nekouzadeh A, *Silva JR*, Rudy Y. "Modeling subunit cooperativity in opening of tetrameric ion channels", *Biophysical Journal*. 2008 October;95(7):3510-20. [PMID: 18621838]
- 59. Rudy Y and *Silva JR*. "Computational biology in the study of cardiac ion channels and cell electrophysiology." *Quarterly Reviews in Biophysics*. 2006 Feb;39(1):57-116. [PMID: 16848931]
- Faber G, Silva JR, Livshitz L, Rudy Y. "Kinetic properties of the cardiac L-type Ca<sup>2+</sup> channel and its role in myocyte electrophysiology: a theoretical investigation." Biophysical Journal. 2007 Mar;92(5):1522-43. [PMID: 17158566]
- 61. *Silva JR* and Rudy Y. "Subunit Interaction Determines I<sub>Ks</sub> Participation in Cardiac Repolarization and Repolarization Reserve", *Circulation*. 2005;112:1384-1391.
- 62. *Silva JR* and Rudy Y. "Mechanism of pacemaking in I(K1)-downregulated myocytes." *Circulation Research*. 2003 Feb 21;92(3):261-3.

#### **Chapters**

- 1. Mangold K, *Silva JR*, "Modeling the Molecular Details of Ion Channels", *Modeling and Simulating Cardiac Electrical Activity*, Ed. Krogh-Madsen, T and Christini, D, 2022.
- 2. Moreno JD, Kang PW, *Silva JR*, "Connecting Cardiac Excitation to the Atomic Interactions of Ion Channels", *Cardiac Electrophysiology: From Cell to Bedside 8<sup>th</sup> ed.* Eds Zipes, Jalife, Stevenson, 2021.
- 3. Cox K, Privitera MB, Alden T, *Silva JR*, Avari Silva JN. "Augmented Reality in Medical Devices". *Applied Human Factors in Medical Device Design*. Academic Press, pp. 327-337, 2019.
- Silva JR. "Slow inactivation of Na<sup>+</sup> channel", Voltage Gated Na<sub>+</sub> Channels, Ed. Peter Ruben, Springer, Berlin Heidelberg, 2014.
- 5. Zhang HX, *Silva JR* and Nichols C. "Cardiac K<sub>ATP</sub> Channels in Health and Diseases", *Electrical Diseases of the Heart, 2nd edition.* pp 259-279, Ed. Ihor Gussak et al., Springer London, 2013.
- 6. *Silva JR* and Rudy Y. "Voltage-Gated Channels and the Heart", *Molecular Machines*. Ed. B. Roux. World Scientific 2011.

7. **Silva JR** and Rudy Y. "Ionic Mechanisms of Ventricular Action Potential Excitation" Cardiac Electrophysiology: From Cell to Bedside, 5th edition, Eds. D.P. Zipes and J. Jalife. Elsevier Science, 2008.

# **Patents and Licenses**

- 1. Silva JNA, **Silva JR**, Southworth MK, Soriano I, Dagley M, Andrews CM, Razzaque S, Hougard I, Tas B, Two-way Communication Between Head-Mounted Display and Electroanatomic System, US Patent App 63/333,900 Provisional.
- Southworth MK, Andrews CM, Soriano I, Henry AB, Silva JR, Silva JN, inventors; Excera Inc, assignee. Multiscale ultrasound tracking and display. United States patent application US 17/740,191. 2022 Nov 10.
  - a. Pending in Europe, Japan, Canada, China
- 3. **Silva JR**, Andrews C, Silva JNAS, Iqbal Z, inventors; Washington University in St Louis WUSTL, assignee. System and method for augmented reality data interaction for ultrasound imaging. United States patent application US 17/799,826. 2023 Mar 2. Licensed to SentiAR/Excera Inc.
  - a. Pending in Korea, Europe, India, Australia
- 4. Southworth MK, Blume W, Silva JN, **Silva JR**, inventors; Sentiar Inc, assignee. Disposable sticker within augmented reality environment. United States patent US 10,733,960. 2020 Aug 4.
  - a. Continuation United States patent US 10,964,291. 2021 Mar 30.
  - b. Issued in Japan (7105996), Pending in Hong Kong, Europe, Canada, Australia
- 5. Blume W, Southworth MK, Silva JN, **Silva JR**. Gaze based interface for augmented reality environment. United States patent US 11,199,898. 2021 Dec 14.
  - a. Continuation US Patent App. 17/524,981, 2022
  - b. Issued in Japan (7213899), Pending in Korea, Israel, Europe, Canada, Australia.
- 6. Blume W, Southworth MK, **Silva JR**, Silva JN, Soriano I, inventors; Sentiar Inc, assignee. Electrogram Annotation System. United States patent application US 17/078,889. 2021 Apr 29.
  - a. Continuation US Patent App. 17/940,564, 2022 Allowed on 3/16/2023
  - b. Pending in Australia, Canada, Europ, Hong Kong, Israel
- 7. **Silva JR**, Silva JNA. System and method for virtual reality data integration and visualization for 3D imaging and instrument position data. United States patent US 10,258,426. 2019 Apr 16. Licensed to SentiAR, Inc.,
  - a. Divisional US Patent Application, 16/386,156
  - b. Issued in Hong Kong (HK1263061) and China (CN105395252A), Pending in Europe, Canada
- 8. **Silva JR**, Zhu W, Priori S, Mazzanti A, Naegle K "Predicting response to sodium channel blockers", US Patent App 16/708,312, Provisional.

## **Invited Lectures**

- 1. University of California Davis, Davis, CA, 2024
- 2. The Johns Hopkins University, Baltimore, MD, 2024
- 3. WWNav-conference, Grindelwald, Switzerland, 2024
- 4. 2023-24 Pioneers in Biomedical Research Seminar Program, Virginia Tech, Roanoke, VA, 2023
- 5. Nanion Webinar talk, 2023
- 6. 8th World Congress of Pediatric Cardiology and Cardiac Surgery, Washington DC, 2023

- 7. Heart Rhythm Society Annual Meeting, New Orleans, LA 2023
- 8. Cardiac Electrophysiology Society, New Orleans, LA, 2023
- 9. Nanion User Meeting, Boston, MA, 2023
- 10. Center for Engineering MechanoBiology Colloquium, St. Louis, 2022
- 11. SPECTRA Conference, St. Louis, 2022
- 12. Northwestern University, Department of Pharmacology, Chicago 2022
- 13. FASEB SRC Ion Channel Regulation, Halifax, Nova Scotia, Canada 2022
- 14. Heart Rhythm Society Annual Meeting, San Francisco, CA 2022
- 15. Society for Mathematical Biology, 2021
- 16. American Heart Association Scientific Sessions, 2021
- 17. Washington University Biomedical Engineering Seminar, St. Louis MO 2021
- 18. Cardiology Grand Rounds, Washington University School of Medicine, St. Louis MO 2020
- 19. Venture Café, St. Louis MO 2020
- 20. UC Davis Cardiovascular Symposium, Davis, CA2020
- 21. Channelopathy 2020, Quebec City, (conference postponed)
- 22. Ohio State University, Department of Biomedical Engineering 2019
- 23. Skandalaris Center for Interdisciplinary Innovation and Entrepreneurship, 2019
- 24. Department of Biochemistry and Molecular Biophysics, Washington Univ School of Medicine, 2019
- 25. University of Michigan, Ann Arbor, Michigan, 2018
- 26. CECAM Multiscale Modeling Workshop, Lugano, Switzerland, 2018
- 27. St. Louis University Biomedical Engineering Society, St. Louis, USA, 2018
- 28. Center for Cardiovascular Research, Washington University School of Medicine, St. Louis USA, 2018
- 29. Cardiac Arrhythmia Mechanisms Gordon Research Conference, Ventura, California, USA, 2017
- 30. Experimental Biology Meeting, Chicago, IL, USA, 2017
- 31. Biomedical Engineering Seminar, Washington University in St. Louis, St. Louis, MO, USA, 2016
- 32. Cardiac Bioelectricity and Arrhythmia Center 10<sup>th</sup> Anniversary Symposium, Washington University in St. Louis, St. Louis, MO, USA, 2015
- 33. Center for Cardiovascular Research, Cardiovascular Research Day, Junior Faculty Presentation, Washington University in St. Louis, St. Louis, MO, USA, 2014
- 34. Biochemistry and Molecular Biophysics Chalk Talk, Washington University in St. Louis, St. Louis, MO, USA, 2014
- 35. Medical Scientist Training Program MAP Talk, Washington University in St. Louis, St. Louis, MO, USA, 2014
- 36. Cardiac Bioelectricity and Arrhythmia Center Seminar, Washington University in St. Louis, St. Louis, MO, USA, 2014
- 37. Université de Montréal (GEPROM), Montréal, Québec, Canada, 2013
- 38. Biophysical Evenings, Washington University in St. Louis, St. Louis, MO, USA, 2013
- 39. University of British Columbia, Vancouver, British Columbia, Canada, 2012
- 40. Simon Fraser University, Vancouver, British Columbia, Canada, 2012

# **Teaching**

BME 440 Introductory Python with Biomedical Applications

Role: Co-Instructor

Year Taught: 2021, 2022, 2023 Avg. Enrolled Students: 40 undegrad

BME 301 Quantitative Physiology

Role: Course Master (bi-annual) Avg. Enrolled Students: 65 undergrad

Year(s) Taught: 2018, 2019, 2020, 2021, 2022, 2023

BME 401 Senior Design

Role: Team taught (3) Avg. Enrolled Students: 77 undergrad

Years taught: 2013, 2014, 2015, 2016

BME 450/550 Numerical Methods for Computational Modeling in Biomedicine (Tier 1 Class)

Role: Developed, Instructor Avg. Enrolled Students: 19 (25% grad, 75% undergrad)

Years taught: 2014, 2015, 2016, 2017, 2018

BME 5901 Integrative Cardiac Electrophysiology

Role: Team-taught (3)

Avg. Enrolled Students: 4

Years taught: 2014, 2016, 2018

## **Trainees**

#### Graduate

Current PhD:

Martina Marras (2021-)

Thesis Topic: Novel Approaches for Class I Anti-arrhythmic Therapies

Current status: Transitioning from MS program to PhD

Chang Hi Lee (2020-)

Thesis Topic: Using Machine Learning Algorithms to Improve Anti-arrhythmic Therapy

Current status: Transitioning from MS program to PhD

Druv Bhagavan (2020-)

Thesis Topic: Algorithms to connect micro-heart muscle phenotype to patient arrhythmia

Anticipated Graduation: 2023

Lucy Woodbury (2020-)

Thesis Topic: Regulation of Nav1.5 by intracellular fibroblast growth factors

Anticipated graduation: 2024

Emily Wagner (2017 - )

Thesis Topic: Disordered protein regulation of Na<sub>V</sub>1.5

Anticipated graduation: 2022

Past PhD:

Wandi Zhu (2013 - 2018)

Thesis topic: Modulation of Cardiac Na<sup>+</sup> Channel Gating by Interacting Molecules

Currently a postdoctoral fellow at Harvard Medical School

Kathryn Mangold (2016 - 2021)

Thesis topic: Computational modeling of cardiac Na<sup>+</sup> channel gating

Currently a post-doctoral trainee at Mayo Clinic

Po Wei Kang (2019 -2021)

Thesis Topic: Calmodulin regulation of cardiac ion channels.

Currently completing medical school.

Paweorn Angsutararux (2017 - 2022)

Thesis topic: Novel methods to monitor ion channel gating in myocytes

Planning to start postdoctoral training at UC Davis

Past MS:

Nicholas Bach (2020 - 2021)

Thesis topic: Tracking algorithm for otolaryngology procedures

Currently working in the lab.

### Post-graduate

Current:

Heather Struckman, PhD, (2023-)

Past:

Jonathan Moreno MD, PhD (2017 - 2021), Assistant Professor, Washington University Khrystyna Sukanova, PhD (2021 - 2022), Research Scientist, Washington University Christopher Andrews, PhD (2018-2020), Sr. Systems Engineer, SentiAR Inc. Divya Kernik, PhD (2019 -2020), Medical Writer, Ashfield MedComms Kuo-Chan Weng, PhD (2017 - 2020), Director iPSC Core, Washington University Zoltan Varga, PhD (2013-2014), Asst. Professor, Debrecen Medical School, Hungary Darya Urusova, PhD (2014-2015), Staff Scientist, Washington University School of Medicine Arie Krumholz, PhD (2014-2015), Data Scientist,

## **Undergraduate (>6 months)**

Past:

Irene Zhou (2021-2022), Graduate Student at Boston University

Anish Bedi (2019-2022), Applying to Medical School

Namit Sambare (2020-2022), Applying to Medical School

Andrew Whitaker (2019-2020), Graduate Student, Washington University

Michael Rudokas (2012-2013), Postdoctoral Fellow, Yale University

Bicong Li (2015-2017), Medical Student

Eric Hsu (2013-2015), MD/ PhD Candidate UT Southwestern

Zach Teed (2014-2017) - PhD Candidate, Princeton University

Chen Zhao (2013-2014) – PhD Candidate, The Johns Hopkins University

## **Professional Associations**

#### Member

2018 - Present. American Heart Association

2016 – Present. Biomedical Engineering Society

2015 – Present. Society of Hispanic Professional Engineers

2015 – Present. Society of General Physiologists.

2011 - Present. Heart Rhythm Society

2010 - Present. American Heart Association

2009 – Present. Biophysical Society

2007 - Present. AAAS/Science

# **University and Professional Leadership**

#### **Department**

Director of Diversity (2020-)

Undergraduate Studies Committee (2013-2014, 2017-2020)

Graduate Studies Committee (2014-2016, Cardiovascular Engineering Program Leader)

Faculty Search Committee (2014, 2016, 2019, 2021)

Masters Studies Committee (2016-2019)

#### School

Advancing Data Science for Racial Equity Faculty Recruiting Committee

Faculty Advisor, Society of Hispanic Professional Engineers, Undergraduate Chapter

Representative to Society for the Advancement of Chicanos/Hispanics and Native Americans in Science

Member, Engineering Space Committee

Panel, Society of Hispanic Professional Engineers, Q&A

Panel, Graduate School Forum

Panel, Engineering Visit Day

## University

Member, Race and Ethnicity Hiring Initiative, 2020-present

Member, University Public Safety Committee, 2020-2021

Member, Conflicts of Interest Committee, 2018-present

Oversight sub-committee

Member, Danforth COVID-19 Re-opening Committee, 2020

Member, HHMI Gilliam Fellowship Nomination Review Committee, 2019-2020

Member, Ethnicity Cluster Hire Review Committee, 2020

Reviewer, SURF Undergraduate Research Fellowship

Participant, Libraries Data Curation Pilot

Participant, StemFIT Program

#### **External**

#### **Grant Review**

National Institutes of Health, MPPB Study Section, Standing Member (2021-2026)

National Institutes of Health, ESTA Study Section, Standing Member (2020-2021)

American Heart Association, Career Development Award (2020)

American Heart Association, Strategically Focused Research Network, Sudden Cardiac Death (2019)

National Institutes of Health, ESTA Study Section, Ad Hoc (2018)

American Heart Association, Transformational Project Award (2018-2019)

American Heart Association, Strategically Focused Research Network, Atrial Fibrillation (2018)

American Heart Association, Electrophysiology 1, permanent member (2015 - 2017)

American Heart Association, Biomedical Engineering Basic Science 3, ad hoc reviewer (2x)

FWO, Belgian Research Foundation, Ad hoc reviewer (4x)

Isaac Newton Trust, Ad hoc Reviewer

Israel Science Foundation, Ad hoc Reviewer

**Arch Grants Pitch Competition** 

#### Oversight

American Heart Association, SFRN: Atrial Fibrillation Oversight Committee (2018 -)

Heart Rhythm Society Journal Subcommittee

#### **Outreach & Diversity**

Brightpath Middle School Outreach Program Host Lab

Council of Diversity for Biomedical Engineering

Aim High St. Louis Career Day

Mentor, Sister Thea Bowman School, Under-represented 8th Grade Students

Invited Speaker, Forsyth School 2<sup>nd</sup> Grade, Topic: Biomedical Engineering (2016 - 2020)

Coach, Lego League, Forsyth School Judge, Honors Science Fair, St. Louis

## **Journal Affiliations**

#### **Editorial Board:**

Frontiers in Neuroscience

#### Reviewer:

**PLoS ONE** 

**American Physical Society** 

Chaos

MethodsX

Journal of General Physiology American Journal of Physiology

Scientific Reports Circulation Research

Circulation: Arrhythmia and

Electrophysiology Heart Rhythm Journal

Journal of the American Heart Association

**Biophysical Journal** 

**Physical Biology** 

Journal of Medical and Biological

Engineering

Journal of Biomechanical Engineering Journal of Molecular and Cellular

Cardiology

Journal of Neuroscience Cardiovascular Research

Europace

**Progress in Biophysics** 

eLife

# **Center Affiliations**

Center for Cardiovascular Research (CCR)

Center for Science and Engineering of Living Systems (CSELS)

Cardiac Bioelectricity and Arrhythmia Center (CBAC)

Center for the Investigation of Membrane Excitability Diseases (CIMED)

Diabetes Research Center (DRC)