Date September 2023

Name Sergej Djuranovic

Contact Information

Address, Telephone and email:

- Office -
 - 0 314-362-9706
 - Department of Cell Biology and Physiology Washington University School of Medicine 660 South Eucllid Avenue, Campus Box 8228 St. Louis. MO 63110-1093
- Home
 - 0 443-825-5217
 - o 1196 Laven Del Lane St. Louis, MO 63122
- sergei.diuranovic@wustl.edu

Present Position

2023 – present Professor of Cell Biology and Physiology Washington University

School of Medicine in St. Louis

Education (degrees in forward chronological order)

Year; degree; degree granting institution

Undergraduate

1999 BSc Biochemistry, School of Chemistry, University of Belgrade, Serbia

BSc thesis work under supervision of Prof. Dr. Vukic Soskic

"Cloning and expression of the terminal intracellular region of human

dopamine D1 and D5 receptors as fusion proteins in E. coli"

Graduate

2001 MSc Biochemistry, School of Chemistry, University of Belgrade, Serbia

MSc thesis work under supervision of Prof. Dr. Vukic Soskic

"Cloning and expression of selected intercellular regions of the human D1 dopamine receptor and analysis of their interactions with subunits

of GO, GS and Gi proteins"

• Djuranovic S, Predic J, Backovic M, Soskic V, Interaction of intracellular loops of dopamine D1 receptor with G protein subunits, 2002, Yugoslav Med. Biochemistry; 21:255-260 (published part of the thesis:

https://www.researchgate.net/publication/247873727 Interaction o f intracellular loops of dopamine D-1 receptor with G protein subunits)

- 2006 PhD Biochemistry, Eberhard Karls University & Max-Planck Institute for Developmental Biology, Tübingen, Germany PhD thesis work under supervision of Prof. Dr. Andrei Lupas "Evolution of the double-psi-barrel fold. Structure and function of Nterminal domains of the AAA proteins"
 - Djuranovic S, Evolution der Substraterkennungsdomänen von AAA-Proteinen: Evolution of substrate recognition domains of the AAA proteins. PhD thesis 2006 (https://core.ac.uk/download/pdf/56755772.pdf)

Postgraduate

- 2006 2007 Postdoctoral Research Fellow Max-Planck Institute for Developmental Biology, Tübingen, Germany (Andrei Lupas, PhD)
 Biochemistry and Structural Biology
 "Structure and function of 19S subunit of archaeal proteasome"
- 2007 2012 Postdoctoral Research Fellow Howard Hughes Medical Institutes and Johns Hopkins University School of Medicine Baltimore, MD (Rachel Green, PhD)

 Biochemistry and RNA biology

"Mechanisms of gene regulation by microRNAs. Regulation of mRNA translation and decay in eukaryotic cells."

Academic Positions / Employment:

2001 – 2002	Research Associate at Clinical Biochemistry Laboratory Children's Hospital at Clinical Center of Montenegro, Montenegro
2013 - 2019	Assistant Professor of Cell Biology and Physiology Washington University School of Medicine, St. Louis, MO
2019 -2023	Associate Professor (with tenure) of Cell Biology and Physiology Washington University School of Medicine in St. Louis

<u>University and Hospital Appointments and Committees</u>

2013 – present	PhD candidate interviewer for MSTP; Molecular Cell Biology program; Biochemistry, Biophysics, and Structural Biology program; Developmental, Regenerative and Stem Cell Biology program; Molecular Genetics and Genomics program and Computational and Systems Biology program
2013 - 2015	Organizer of Cell Biology and Physiology Departmental Retreat
2013 – present	Member of PhD Admissions Committee for Molecular Cell Biology program
2016 - present	Member of PhD Admissions Committee for Biochemistry, Biophysics, and Structural Biology program
2016 – 2022	Member of Cell Biology and Physiology Faculty Search Committee (successful recruitment of new faculty members Drs. Silvia Jansen, David Kast and Amber N Stratman)
2017 – present	Institute of Clinical and Translational Sciences liaison for Cell Biology and Physiology Department
2018 - 2023	Chairman of PhD Admissions Committee for Molecular Cell Biology program
2021 - 2023	Member of Strategic planning for Molecular Cell Biology PhD program and CBP Department
2022 – present	Siteman Cancer Center Health Equity Fellow, faculty for community outreach, community engagement, and health disparities. Program for the Elimination of Cancer Disparities (PECaD), advocating for the inclusion of community-engaged opportunities and research.
2022 – present	Advisory Board member of Siteman Cancer Center Diversity, Equity, and Inclusion Committee
2023 – present	Vice-Chair Elect for the Executive Committee of the Faculty Council

Honors and Awards

1994-1999	National Fellowship for Talented Students in Sciences and Arts, Ministry of Science, Montenegro
2002-2006	Doctoral Fellowship, Max-Planck Society, Germany
2006	Award for best presentation, American Society for Biochemistry and Molecular Biology, 2006 ASBMB/JBC centennial meeting, San Francisco, USA
2006	Summa Cum Laude, PhD with honors, Eberhard Karls University and Max-Planck Institute for Developmental Biology, Tübingen, Germany
2007-2012	Postdoctoral Fellowship, Howard Hughes Medical Institute & Johns Hopkins University School of Medicine, Baltimore, USA
2013-2019	Elected member of Center for Young Scientists, Montenegrin Academy of Science, Montenegro
2014	Organizer of Cell Biology and Physiology Retreat, Rend Lake Resort in Whittington, Illinois
2016	Invited speaker and lecturer, Ministry of Science of Montenegro, Summer School of Science, Podgorica, Montenegro
2017	Invited panelist, "Links between science and economy", South European and European Union Forum, Budva, Montenegro
2017	Children Discovery Institute and Zebrafish facility Award
2017	LEAP Inventor Challenge Award, Skandalaris Center and Washington University, St. Louis
2017	Elected representative of Serbian Biochemical Society for FEBS (The Federation of European Biochemical Societies)
2018	Finalist of the Burroughs Wellcome Fund Investigators in the Pathogenesis of Infectious Disease Award (PATH)
2018	Keynote speaker, Systems Biology Conference and workshop, St. Petersburg, Russia
2018	SBS plenary speaker and chair, From Molecules to Organisms meeting, FEBS 3+, Siofok, Lake Balaton, Hungary
2018	Session Chair mRNA turnover, Translation Control meeting, Cold Spring Harbor, USA

2019	LEAP Inventor Challenge Award, Skandalaris Center and Washington University, St. Louis
2019	Invited reviewer for Siteman Cancer Center
2019	Reviewer for NSF Gene expression grants, USA
2019	Reviewer for Wellcome Trust Grants, UK
2019	Member of Centre of Excellence for Biomedical Research in Montenegro
2020	Keynote speaker, Systems Biology Conference and Workshop, Melbourne, Australia
2020	Invited member of NIGMS Established Investigator R35/MIRA study section
2020	Awardee of Chan Zuckerberg Initiative for Collaborative Pair Pilot Project Awards on Neurodegeneration
2021	Siteman Investment Award
2021	Permanent member of NIH/NIGMS Molecular Genetics B Study Section (MGB)
2021	Permanent member of NIH/NIGMS Molecular Genetics Study Section (MG)
2022	Member of the Health Equity Fellowship within Siteman Cancer Center and the Program for the Elimination of Cancer Disparities (PECaD), advocate for inclusion of community-engaged opportunities and research.
2022	Chairman of the reviewing committee for Immunology Graduate program at DBBS Washington University
2022	Advisory Board member Siteman Cancer Center Internal Diversity, Equity, and Inclusion Committee
2022	Member of scientific advisory board of rBIO Biotech Company
2022	Co-chair for MG study section at NIGMS/NIH
2022	Awardee of Chan Zuckerberg Initiative for Collaborative Pair Awards on Neurodegeneration

2022	Reviewing editor for Frontiers in RNA Research
2023	Scientific reviewer for MRC Toxicology Unit Department at the University of Cambridge, UK
2023	Editorial Board Member Journal of Biological Chemistry

Editorial Responsibilities

Editorial board member

Journal of Biological Chemistry

Reviewing editor

Frontiers in RNA Research – section for non-coding RNA

<u>Reviewer</u>

Nature

Science

Nature Communications

Nature Structural and Molecular Biology

Cell

Molecular Cell

Nucleic Acid Research

eLIfe

Cell Reports

Cell Systems

Current Biology

Journal of Biological Chemistry

Scientific Reports

Cell Proliferation

RNA journal

WIRE's RNA

Plos One

Plos Pathogens

Plos Genetics

Cellular and Molecular Biology Letters

ACS Chemical Biology

Journal of Cellular and Molecular Medicine

Invited guest editor

Methods in RNA (declined) WIRE's RNA (declined)

National Scientific Panels

2017	Links between science and economy", South European and European Union Forum, Budva, Montenegro
2017	Evaluator for collaborative scientific projects between European Union and Montenegro
2018	Expert panel for Accreditation and Higher Education Institutions Reaccreditation, Government of Montenegro
2019	Member of reviewing panel for Siteman Cancer Center
2019/2022	Reviewer for NSF Gene expression grants, USA
2020	Member of NIGMS Established Investigator R35/MIRA study section
2021	Permanent member of NIH/NIGMS Molecular Genetics B (MGB) and Molecular Genetics (MG) Study Section.
2023	Member of NSF-MCB Genetic Mechanisms review panel

Community Service Contributions (year-year of service)

- PhD candidate interviewer for MSTP; Molecular Cell Biology program; Biochemistry, Biophysics, and Structural Biology program; Developmental, Regenerative and Stem Cell Biology program; Molecular Genetics and Genomics program and Computational and Systems Biology program
- Organizer of the Cell Biology and Physiology Departmental Retreat
- Member of PhD Admissions Committee for Molecular Cell Biology program
- Member of PhD Admissions Committee for Biochemistry, Biophysics, and Structural Biology program
- Member of Cell Biology and Physiology Faculty Search Committee (successful recruitment of new faculty members Drs. Silvia Jansen, David J. Kast and Amber N. Stratman)

- Institute of Clinical and Translational Sciences liaison for Cell Biology and Physiology Department
- Chairman of PhD Admissions Committee for Molecular Cell Biology program
- Member of Curriculum Committee for Division of Biomedical and Biological Science (DBBS) programs at Washington University in St. Louis
- Member of the Health Equity Fellowship within Siteman Cancer Center and the Program for the Elimination of Cancer Disparities (PECaD), advocate for inclusion of community-engaged opportunities and research.
- Advisory Board member for Siteman Cancer Center Internal Diversity, Equity, and Inclusion Committee
- Vice-Chair Elect for the Executive Committee of the Faculty Council

Professional Societies and Organizations

- Member of Serbian Biochemical Society
- Member of The American Society for Biochemistry and Molecular Biology (ASBMB)
- Center for Young Scientists, Montenegrin Academy of Science, Montenegro
- Member of RNA Society
- Siteman Cancer Center, School of Medicine, Washington University in St. Louis
- Member of the American Association for Cancer Research (AACR)
- Representative of Serbian Biochemical Society in FEBS
- Hope Center, School of Medicine, Washington University in St. Louis
- Member of Centre of Excellence for Biomedical Researches (CEBIMER) of Montenegro
- Member of American Society of Gene & Cell Therapy (ASGCT)
- Member of American Society for Microbiology

Media activities

On Argonaute miRNA studies

https://scienceblog.com/29634/argonautes-a-big-turn-off-for-proteins/

https://www.hhmi.org/news/researchers-reveal-new-details-about-how-argonaute helps-halt-protein-production

On translation ramp and protein synthesis

https://www.labmanager.com/news/scientists-find-way-to-supercharge-protein-production-21281

https://phys.org/news/2019-12-scientists-supercharge-protein-production.html

https://www.genengnews.com/news/in-the-hurdle-race-to-boost-protein-production-dont-trip-over-codons-3-5/

 $\frac{https://www.outlookindia.com/newsscroll/new-method-ramps-up-protein-production-by-thousandfold-study/1690184$

https://www.business-standard.com/article/pti-stories/new-method-ramps-up-protein-production-by-thousandfold-study-119122000252 1.html

https://m.dailyhunt.in/news/india/english/indiatvnews-epaperintvnews/new+method+ramps+up+protein+production+by+thousandfold-newsidn154379074

 $\frac{https://health.economic times.indiatimes.com/news/diagnostics/new-method-ramps-up-protein-production-by-thousand fold/72897699$

https://bioengineer.org/scientists-find-way-to-supercharge-protein-production/

https://www.techexplorist.com/supercharging-protein-production/28619/?utm_source=rss&utm_medium=rss&utm_campaign=supercharging-protein-production

https://6park.news/washington/hookworms-have-potential-to-protect-soldiers-from-chemical-and-biological-weapons-washington-university-school-of-medicine-in-st-louis.html

On creation of hypomorphic mutants using polyA tracts

https://coboscientific.com/genome-editing/tunr-flexible-gene-editing-system/

https://techstartups.com/2018/05/22/gene-editing-startup-canopy-biosciences-raises-2-4-million-series-financing-advance-development-research-tools/

https://medicine.wustl.edu/news/new-genetic-engineering-technique-help-designstudy-biological-systems/

https://www.scientifictechnologynews.com/canopy-biosciences-licenses-novel-gene-editing-technology-washington-university-johns-hopkins-university/

https://canopybiosciences.com/our-news/page/3/

https://www.eurekalert.org/multimedia/pub/131133.php

 $\frac{https://www.prnewswire.com/news-releases/canopy-biosciences-licenses-novel-gene-editing-technology-from-washington-university-and-johns-hopkins-university-300432392.html$

https://www.biotechniques.com/bioengineering-biophysics/a-faster-way-to-generate-hypomorphic-mutations/

https://igtrcn.org/rapid-generation-of-hypomorphic-mutations/

https://web.genewiz.com/webinar hypomorphic alleles

https://www.sciencedaily.com/releases/2017/01/170120091004.htm

http://www.mariakonovalenko.com/blog/2017/1/23/researchers-invent-a-method-to-produce-a-precise-amount-of-protein-on-demand

https://gmoanswers.com/studies/study-new-genetic-engineering-technique-could-help-design-study-biological-systems

http://www.frontlinegenomics.com/news/10901/canopy-biosciences-acquire-rights-gene-editing-technology/

On polyA research

https://www.the-scientist.com/daily-news/aaaaa-is-for-arrested-translation-35106

<u>ASBMB</u>

 $\frac{http://www.asbmb.org/uploadedFiles/ASBMBToday/Content/Archive/ASBMBToday-2006-7.pdf}{}$

In Serbo-Croatian language

http://www.rtcg.me/vijesti/ekonomija/182757/regionalna-konferencija-o-ekonomiji-u-budvi.html

http://www.mneresearch.ac.me/cg/vijest.php?id=64

Major Invited Professorships and Lectureships

2005	Invited speaker for colloquium of the MPI for Developmental Biology and Friedrich Miescher Laboratory, Tuebinge, Germany
2006	Speaker for ASBMB Annual Meeting, San Francisco, USA
2011	Invited speaker for translational control interest meeting, Johns Hopkins University School of Medicine, Baltimore, USA
2011	Speaker at Protein Synthesis and Translational Control Conference, EMBL Heidelberg, Germany
2012	Invited speaker at Department of Biochemistry, School of Medicine, University of Utah, Salt Lake City, USA
2012	Invited speaker at Department of Biology, Johns Hopkins University, Baltimore, USA
2012	Invited speaker at Department of Cell Biology and Physiology, School of Medicine, Washington University, St. Louis
2012	Invited speaker for Symposium on Systems Biology at Max Delbruck Center for Molecular Medicine, Berlin, Germany

2012	Invited speaker and lecturer, Belgrade Food and Allergy International Conference, Horizons in Life Sciences, FCUB ERA, Belgrade, Serbia
2012	HHMI Science Meeting, Janelia Research Campus, Ashburn, VA
2013	Invited lecture at Siteman Cancer Center – Breast Cancer Research Program, St. Louis, USA
2013	Invited lecture at Department of Genetics, School of Medicine, Washington University, St. Louis, USA
2014	Invited lecture at Department of Molecular Microbiology, School of Medicine, Washington University, St. Louis, USA
2014	Invited speaker and lecturer, Prona Summer School of Science, Cetinje, Montenegro
2014	Speaker at Translational Control Meeting, Cold Spring Harbor, USA
2015	Invited lecture at Department of Biology, Western Ontario University, London, Ontario, Canada
2015	Invited speaker at Eukaryotic mRNA processing meeting, Cold Spring Harbor USA
2015	Invited lecture at Department of Biology, University of Missouri St. Louis, USA
2016	Invited lecture at Department of Developmental Biology, School of Medicine, Washington University, St. Louis, USA
2016	Speaker at Post-Transcriptional Gene Regulation, Gordon Research Conference, Stove, VT, USA
2016	Speaker at Translational Control Meeting, Cold Spring Harbor, USA (talk given by PhD student Laura Arthur)
2016	Keynote speaker for Biochemistry and CMB Program Retreat, Department Of Molecular Biophysics and Biochemistry, Washington University, New Heaven, MO, USA
2016	Invited Lecture at Department of Ophthalmology, Vision Science seminar, School of Medicine, Washington University, St. Louis, USA
2017	Invited lecture at Breast Cancer Panel Group, Department of Oncology, School of Medicine, Washington University, St. Louis, USA
2017	Speaker at Tropical Medicine and Parasitology Seminar, Department of Microbiology, School of Medicine, Washington University, St. Louis, USA
2017	Speaker at Translation Machinery in Health & Disease, Gordon Research Conference, Galveston, TX, USA (talk given by PhD student Kyle Cottrell)

2017	Speaker at RNA Society Meeting, Prague, Czech Republic (talk given by PhD student Laura Arthur)
2017	Invited lecture at Polish National Academy, Institute of Biophysics and Bioinformatics, Warsaw, Poland
2017	Speaker at Protein Synthesis and Translational Control Meeting, EMBL Heidelberg, Germany
2017	Speaker at Transgenic Technology Meeting, Snowbird resort, Salt Lake City, Utah, USA (talk given by Rachel Delston from Canopy Biosciences)
2017	Invited lecture at RNA Institute, School of Medicine, Case Western University, Cleveland, USA
2017	Invited lecture at Lady Davis Institute, SMBD Jewish General Hospital, McGill University, Montreal, Canada
2017	Invited webinar, Creation of Hypomorphic Alleles for Functional Gene Studies, Genewiz
2018	Keynote lecture, Systems Biology Conference and Workshop, St. Petersburg, Russia
2018	Plenary speaker, Keynote speaker and chair, From Molecules to Organisms Conference, FEBS 3+, Siofok, Lake Balaton, Hungary
2018	Session chair and invited speaker, mRNA turnover session, Translation Control Meeting, Cold Spring Harbor, USA
2019	Invited speaker for MBG Young Investigator Symposium 2019, Department of Molecular Biology and Genetics, Aarhus University, Denmark
2019	Invited lecture at Pharmaceutical Institute, University of Kiel, Dept. of Pharmaceutical and Medicinal Chemistry, Germany
2020	Invited speaker for Systems Biology Workshop at La Trobe University, Melbourne, Victoria, Australia
2020	Invited lecture Saint Louis University, Department of Biochemistry, Saint Louis, USA
2020	Invited Lecture UT Health San Antonio School of Medicine, Department of Biochemistry and Structural Biology, San Antonio, USA
2021	Invited speaker for Systems Biology Workshop at La Trobe University, Melbourne, Victoria, Australia
2022	Invited Speaker UC Riverside Molecular, Cell and Systems Biology Department, Riverside, USA
2022	Speaker at tRNA meeting at Columbus, OH, USA

2022	Invited Speaker for Ribosome Structure and Function meeting in Bordeaux, France
2022	Speaker at Translational Control Meeting, Cold Spring Harbor, USA
2022	Invited lecture at THINC RNA Biology and Discovery Center at Baylor College of Medicine, Houston, USA
2023	Invited lecture at Department of Genetics, School of Medicine, Case Western University, Cleveland, USA
2023	Invited lecture at Department of Oncology at Baylor College of Medicine, Houston, USA
2023	Invited speaker and panelist for RNA Biology session Neurodegeneration Challenge Network (NDCN) at Chan-Zuckerberg Annual Meeting, San Diego, USA
2023	Invited lecture at Fred Hutch Cancer Center and Department of Biochemistry, University of Washington, Seattle, USA
2023	Speaker at Eukaryotic mRNA Processing, Cold Spring Harbor, USA
2023	Invited lecture at Max-Planck Institute for Biology, Tübingen, Germany
2023	Invited lecture at Montenegro Academy of Science , Podgorica, Monetnegro
2023	Invited lecture at XXX Meeting of the Balkan Clinical Laboratory Federation, Herceg Novi, Montenegro
2023	Invited lecture at Johns Hopkins University, School of Medicine, Department of Molecular Biology and Genetics
2024	Invited lecture at RNA Therapeutics Institute at UMass Medical School, Worcester, USA
2024	Invited speaker for RNA Regulation meeting at Stowers Institute for Medical Research in Kansas City, Missouri

Research

<u>Completed Research Support</u>

IRG-58-010-58-2 Djuranovic (PI) Siteman Cancer Center/American Cancer Society

01/01/15-31/12/15

Regulation of Genes by Ribosome Stalling in Cancer Cell Growth The goals of this project are to investigate the notion that ribosome stalling effects gene regulation through polyA-motifs and to examine how deregulation of this pathway leads to tumorigenesis.

Role: PI

LEAP Inventor Challenge Award, Djuranovic (PI)

10/01/17-31/12/19

Fine tuning of gene expression by polyA tracks

Project goal is to de-risk and further develop a technology that fine-tunes gene expression to desirable amounts using engineered poly adenosine tags.

LEAP Inventor Challenge Award Djuranovic (PI)

04/01/19-04/01/21

Use of N-terminal peptides for increase in protein expression.

Project goal is to use N-terminal penta-peptides for optimization of the protein synthesis, stability and folding by use of short N-terminal peptides.

LEAP Inventor Challenge Award Djuranovic (PI)

01/07/19-01/07/21

Use of polybasic peptides and polymers as antimalarial agents.

Project goal is to test and commercialize a new lead substance for the treatment of human malaria.

Sponsored Research Funds (rBIO)

20/05/21-20/11/21

Optimization of the bio pharmaceutically important proteins for expression in various translation systems.

Ongoing Research Support

2R01 GM112824-01A, Djuranovic (PI)

01/07/15-30/09/24

Mechanisms for Modulation of miRNA-Mediated Gene Silencing
The main goal of this project is to define cellular mechanisms used to modulate the process
of miRNA-mediated gene silencing. These studies will ultimately give insight into the
complexity of gene regulation at the level of translation and will be relevant to the better
understanding of alterations in these regulatory mechanisms that may lead to various
pathological states.

1R01GM136823-01A1, Djuranovic (PI)

06/01/21 - 03/31/25

Plasmodium falciparum, a causative agent of human malaria, inflicts an enormous burden on global human health. Even though P. falciparum is a eukaryotic organism, there are vast differences in the mechanisms of mRNA translation and surveillance between P. falciparum and humans. This project will apply new approaches to understand better the evolution of the previously unexplored process of protein synthesis in the AT-rich genome of malaria parasites and exploit new parasite-specific drug targets.

SIP Research Development Awards 823924. Diuranovic (PI)

01/04/21 - 01/04/24

This project follows mutations in polyA track motifs of ZCRB1 and RNPC3 minor spliceosome genes and their impact on gene expression levels of multiple genes involved in tumor suppression and tumorigenesis. These mutations are thought to be silent; however the same mutations lead to gene dosage effects potentially causing tumorigenesis.

1R01 MH116999, Djuranovic (5% Co-PI)

10/01/19-09/30/24

National Institute for Mental Health – NHM

Highly parallel analysis of 5' and 3' UTR variants in Autism Spectrum Disorders Studies on the role of mutations in non-coding regions of mRNA and their role in alleviating mechanisms of gene expression in Autism Spectrum Disorders (PI – Dougherty)

Simons Foundation Autism Research Initiative, Djuranovic (2% Co-PI)

01/10/18-30/09/23 (extended for an additional year)

SSC-ASC Whole Genome Sequencing Consortium: UTR Mutation Analysis Mutation analysis of non-coding mRNA regions in patients with behavioral diseases. (PI – Dougherty)

Children Discovery Institute and Zebrafish facility Award Djuranovic 100%

12/01/16-12/2023

Ribosome stalling and frameshifting in zebrafish model.

Role of polyA track motifs, NGD and NMD pathways in the regulation of ribosome biogenesis during the early zebrafish embryogenesis.

DoD/DARPA/Charles River Analytics Co-PI 8% Djuranovic) 15/04/2021-14/04/2026

Personal Protective Biosystems – Membranes and Commensal Helminths (MaCH) Vectored prophylaxis and neutralization of the program specified chemical/biological (CB) agents. Gene optimization toolkit contribution of Djuranovic Lab.

Sponsored Research Funds (rBIO)

08/03/22-12/09/23

Optimization of the bio pharmaceutically important proteins for expression in various translation systems.

Chen Zuckerberg Initiative (50% PI Djuranovic) phase I	2020-2022
Chen Zuckerberg Initiative (50% PI Djuranovic) phase II	2022-2026

Increasing mRNA translation to treat neurodegeneration

To develop universal gene-targeting strategies that stimulate mRNA translation to understand and treat neurodegenerative diseases.

RTW Charitable Foundation (co-PI with Dougherty, Kroll and Miller) 04/2024

Develop ASOs to restore normal MYT1L levels

Short term goals are to: (1) test the ability of in vivo massively parallel reporter assay technologies (MPRAs) to predict ASO targetable elements, and (2) identify a lead ASO compound for preclinical rescue studies. The long term goals are to conduct preclinical rescue studies in mouse and human models. The long term objectives of these goals are both to enable future clinical trials or n=1 studies for MYT1L Syndrome with the ASO developed here, and to develop a generalizable pipeline for identifying ASOs for any haploinsufficiency syndrome.

Pending and planned Research Support

Commercial Solutions Proposal DARPA

01/06/23-31/05/24

Hookworm as drug production and delivery technology

Project for commercialization of DARPA grant covered technology

HHMI 2024 Investigator Competition

Regulation and systems biology of protein synthesis

R01GM141288, Djuranovic (PI)

01/06/23-31/05/27

NIH-NIGMS MSFC study section (33.0 percentile first submission)

Structural and functional studies of early elongation events in protein synthesis

The amount of newly synthesized protein is generally defined by rates of translation initiation and elongation. In this proposal we seek to understand the significance of starting penta-peptide sequences in the regulation of protein synthesis. The results of our work will give a comprehensive understanding of the mechanics of early elongation events and the degree to which the first five amino acids of a polypeptide chain and ribosome structure determine the output of protein synthesis.

NICHD R01 planned submission

01/07/23-31/04/27

Continuation of the project on polyA track motifs of ZCRB1 and RNPC3 minor spliceosome genes and their impact on gene expression levels of multiple genes involved in tumor suppression and tumorigenesis.

Sponsored research agreement (Perfect Day Company)

Optimization of the diary product related important proteins for expression in various translation systems.

Trainee/Mentee/Sponsorship Record:

Mentored PhD thesis (defended)

- 2013 2017 Dr. Kyle A Cottrell, PhD thesis title "Regulation of gene expression by RNA binding proteins and microRNAs", successfully defended December 2017, postdoctoral work at the lab of Dr. Jason Weber
- 2014 2017 Dr. Laura L Arthur, PhD thesis title "Mechanism of Gene Regulation by Coding PolyA Tracks", successfully defended September 2017, Stuff scientist in the lab of Dr. Maxim Artyomov
- 2015 2020 Jessey Erath, PhD thesis title "Evolution of Ribosomes in an AU-Rich Transcriptome: Translational Tolerance of Poly(A) Motifs in P. falciparum." postdoctoral work in the lab of Dr. Peng Yuan
- 2017 2023 Geralle Powell, PhD thesis title "Characterizing the significance of polyA tracks in gene regulation and disease". Postdoctoral work at WUSM.

2019 – present Courtney Jungers, proposed PhD thesis title "Modulation of

miRNA-mediated gene silencing by AU-rich binding proteins"

Rotation students

Kyle Cottrell - Lab of Djuranovic, Sergej Molecular Cell Biology

Laura Arthur - Lab of Djuranovic, Sergej Molecular Genetics and Genomics

John Dean - Lab of Lodhi, Irfan Molecular Cell Biology Yansel Nunez - Lab of Pike, Linda Molecular Cell Biology

Jessey Erath - Lab of Djuranovic, Sergej Molecular Microbiology and Microbial

Pathogenesis

Kellan Weston - Lab of Yi, Jason Molecular Cell Biology

Geralle Powell - Lab of Djuranovic, Sergej
Ellie Wilson - Lab of Spencer, David
Molecular Genetics and Genomics
Molecular Genetics and Genomics

Cortney Jungers – Lab of Djuranovic, Sergej Molecular Cell Biology

Jade Enright – Lab of
Katherine Floyd – Lab of Daniel Goldberg
Sarah Koester – Lab of Joe Dougherty

Molecular Cell Biology
Molecular Cell Biology

Klaudio Fatmiri –lab of Andrea Sorrano
Tessa Lochetto - lab of Natali Niemi
Biochemistry and Biophysics
Molecular Cell Biology

Kaylee O'Donell – lab of Katherine Weilbaecher Molecular Cell Biology

<u>Undergraduate Bio200/Bio500 program students</u>

Felipe Bertazzo (Successfully enrolled in Sao Paolo School of Dentistry)
Denise Rogers (Successfully enrolled in JHU School of Public Health)
Manasvi Verma (Currently in PhD program at Harvard University)

Natasha Recoder Roshan (Currently in PhD program at Cornell School of Medicine)

Sivakumar (Currently in empower through health NGO Uganda)

Miles George (Currently in the MIT Biology program)

Eylul Horozoglu (Currently in the WUSTL Genetics PhD program)

Ethan Tsao (Currently in the UCSF medical school)

Danielle Kemper (Currently in Djuranovic Lab)

Patents and Publications

• U.S. and international Patent, Serial No. PCT/US2017/041766 (converted to the full patent in 2017). "Gene Regulation and Rapid Generation of Hypomorphic Mutations Using PolyATracks." Licensed to Canopy Biosciences.

- U.S. and International Patent Application Serial No. 62/540,897 (conversion to the full patent in August 2019, #016819/US-2) "Short N-Terminal Peptides for increasing production of recombinant proteins." (licensed to rBIO)
- US Provisional Patent Application Serial No. 62/696,868 (conversion to full patent expected February 2020) "Antimalarial Compositions and Methods of use thereof."
- U.S. Provisional Patent Application, Serial Number pending, "N-Terminal Peptides for Increasing Folding Kinetics, Chromophore Maturation, and Stability of Florescent Proteins with Beta-Barrel Scaffold."
- 2 additional patents on antisense oligonucleotide drugs for neurodegeneration treatment (co-owned with Tim Miller) and technology for treatment of haploinsufficiency related diseases (co-owned with Tim Miller and Joe Dougherty)

<u>Original, peer reviewed articles in refereed journals (* corresponding or senior author, #co-first author)</u>

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