National Aeronautics and Space Administration



## Biological and Physical Sciences

SPACE BIOLOGY PROGRAM

Sharmila Bhattacharya Space Biology Senior Program Scientist

> Presentation at the BPS Advisory Council Meeting Nov 15, 2022



# Agenda

- Overview of the Space Biology program
- Research Focus Thriving in Deep Space
- Artemis I Space Biology Payloads
- Space Biology Lunar Payload (LEIA and PRISM)
- Accomplishments in Open Science
- Recent and Current Space Biology Opportunities
- Recent Science Highlight

# Overview of the Space Biology Program



## **Objectives**

- Discover how biological systems **respond** to the space environment
- Identify the underlying <u>mechanisms</u> and develop models for biological systems in space
- Provide mechanistic understanding to support human health in space
- Promote <u>open science</u> through the GeneLab Data System and Life Sciences Data Archive
- Develop <u>technologies</u> to enable spaceflight research
- Transfer the knowledge and technology of space-based research to **benefit life on Earth**

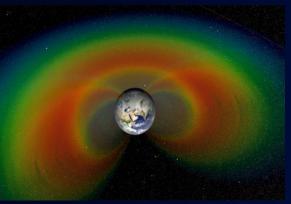
### Biologically Relevant Environmental Factors Encountered in Spaceflight



#### Microgravity/Reduced Gravity

### Ionizing Radiation

#### Altered Day/Night Cycles: Circadian Rhythm Changes



Credits: NASA/Goddard Space Flight Center/Scientific Visualization Studio



Elevated CO2

Reduced atmospheric pressure and elevated volumetric fraction of oxygen Altered Temperature and Atmosphere

Isolation







Regolith

#### COMBINATION OF MULTIPLE STRESSORS

# <u>Thriving in Deep Space (TIDES)</u>

Ground studies
 Space studies
 Ground & space studies

#### Understand Fundamental Mechanisms

Use model organisms to determine how animals/humans respond to deep space environments







Understand how model plants, crops, & seeds respond to and can thrive in deep space environments



#### Build a Foundation for Sustained Life on Mars

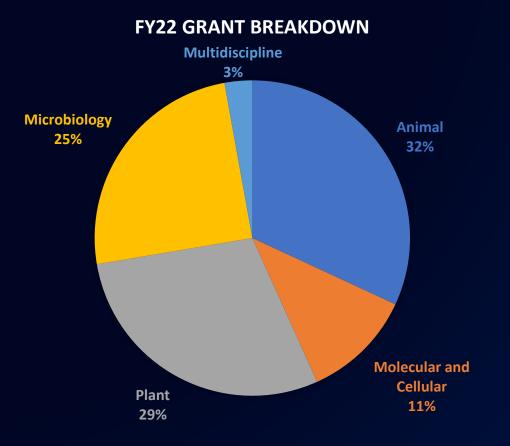
Engineer habitats and ecosystems to enable astronauts' independence from Earth

 Stabilize human/animal-plantmicrobial ecosystems in the context of multiple deep space stressors

#### **DEEP SPACE STRESSORS**

Gravity		R	adiation	Temperature/At	tmosphere	Day/Night/Circadian Cycle			Isolation		
PLATFORM PROGRES	SION										
Ground	Sub	-Orbital	LEO/ISS	Gateway	, ) ι	unar Surface	Mars Transit		Martian Surface		
MODEL ORGANISM P	ROGRESS	ION									
Unicellular (e.g., yea	ost fungi)		brates (e.g., flies, wo	rms, tardigrades)	Vertebr	ates (e.g., mice,		Humans			
Officential (e.g., yea			ole model plants (e.g	., Arabidopsis)	Edible & others (for micronutrients, etc.)				6		

# Space Biology Content



Total SB Grants	141
Flight	62
Ground	79
Flight and Ground	7

Number Directed vs Competed								
Directed	9							
Competed	132							

## Newly Selected Animal Grants: HRP Partner; Radiation emphasis (2022)

### ROSES 2021-E.11: Space Biology: Animal Studies



Insights into the impacts of continuous, low dose-rate neutron radiation exposure on maternal and fetal skeletal physiology Heather Allaway, Ph.D. Louisiana State University and A&M College



Integrated CNS assessment in rodent models of altered gravity and irradiation Joshua Alwood, Ph.D. NASA Ames Research Center



Unraveling the role of mitochondrial dysfunction and senescence on inhibition of tissue regeneration during spaceflight and amelioration by a novel countermeasure, PQQ Elizabeth Blaber, Ph.D. Rensselaer Polytechnic Institute



Feasibility study: Use of neural networks to predict adaptability and multiday performance saving in dual motorcognitive tasks after exposure to space flight stressors Ashley Blackwell, Ph.D. Eastern Virginia Medical School



Characterization of Female Reproductive Health Risks for Long-Duration Spaceflight using Federated Machine Learning Sylvain Costes, NASA Ames Research Center



Sex-specific physiological and transcriptomic CNS responses to combined effects of spaceflight stressors in *Drosophila melanogaster* Janani Iyer, Ph.D. NASA Ames Research Center



Sustained effects of spaceflight on Anemia and severity of effects dependency on age. Cassandra Juran, Ph.D. NASA Ames Research Center



Stressors to Spaceflight: Identification of Transposon-Driven Changes to Gene Networks in GeneLab Data <sup>1</sup> Caralina Marin de Evsikova, Ph.D. Bay Pines Foundation, Inc



Circadian rhythm disruption and gravitational disturbance in a Lunar mission analog: consequences for muscle function during and after the mission Marie Mortreux, Ph.D. Beth Israel Deaconess Medical Center, Inc.



Partial Gravity and Sex-Difference Effects on the Venous Circulation Anand Narayanan, Ph.D. Florida State University



Acute and long-term effects of combined radiation and partial unloading on neurological and musculoskeletal systems in male and female rats Seward Rutkove, Ph.D. Beth Israel Deaconess Medical Center, Inc.



The effect of different genetic mutations and pharmacologic interventions on transcriptional responses to spaceflight *in C. elegans* Craig Willis, Ph.D. Ohio University

## Newly Selected Plant Grants: HRP Partner; Radiation emphasis (2022)

#### ROSES 2021-E.9: Space Biology: Plant Studies



Determining the impact of space radiation and simulated microgravity on plant root microbial community composition and function.

John Baker, Ph.D. Medical College Of Wisconsin, Inc.



How do carbon fixing strategies affect nutritional content under high CO2? A comparison of C3 vs. C4 microgreens? Colleen Doherty, Ph.D. North Carolina State University



Growing Food on Mars: Determining the impact of radiation, atmospheric composition, and rock substrate on plant growth in a Space Rock Garden Experiment Rebecca Lybrand, Ph.D. University Of California, Davis



Plant Trek: Investigating Strategies for Regolith Pre-Conditioning to Support the Establishment of Plant-Microbe Systems in Martian Habitats Kennda Lynch, Ph.D. Universities Space Research Association, Columbia



Temporal Lighting Optimization to Improve Lettuce Productivity and Nutritional Quality Under Superelevated CO2 Stress Qingwu Meng, Ph.D. University Of Delaware



Telomere dynamics and oxidative stress in Arabidopsis in the space radiation environment Dorothy Shippen, Ph.D. Texas A&M AgriLife Research



Leaf Sensor Network for In Situ and Multiparametric Analysis of Crop Stressors Shawana Tabassum, Ph.D. University of Texas, Tyler.

## Space Biology Awardees from ROSES in 2021/2022

#### ROSES-2020 E.12: GeneLab Research Proposals



Alternative Splicing and Transcriptome Modulation under Space Flight Response in Plants Pankaj Jaiswal, Ph.D., Oregon State University, Corvallis, Oregon



In silico Modeling of Gene Networks and Mechanisms Associated with Plant Gravitropism Sushma Naithani, Ph.D., Oregon State University, Corvallis, Oregon



Comparative Analysis of Multi-Gravity Studies on Earth and ISS Sarah Wyatt, Ph.D., Ohio University, Athens, Ohio

#### **ROSES-2020 E.12: Early Career Investigation Proposals**



Responses of Microbes and Microbial Communities to Prolonged Exposure to Space Radiation Jonathan Galazka, Ph.D., NASA Ames Research Center



Responses of Microbes and Microbial Communities to Prolonged Exposure to Space Radiation Jessica Lee, Ph.D., NASA Ames Research Center

These awards are supported in part by the Science Mission Directorate Research Catalyst Fund

## Featured Animal Biology Ground-based Projects

#### ROSES 2020-E.12: Call for Flight and/or Ground Research Proposals



A Multi-omics and multi-species examination of combined environmental stressors of space exploration Dawn Bowles, Ph.D. Duke University, Durham, NC



Develop a Novel Single-Cell Biodosimetry for Brain Genomic Instability and Neurodegeneration to Predict Clinical Health Outcomes in Human Spaceflight Crews Xiaohong Lu, Ph.D, Louisiana State University, Shreveport, Louisiana



Osteocyte Plasma Membrane Disruptions in Skeletal Adaptation to Loading and Unloading Meghan McGee-Lawrence, Ph.D., Augusta University Research Institute, Inc., Augusta, Georgia



Thrombosis in microgravity Anand Ramasubramanian, Ph.D., San Jose State University, San Jose, California



A Technology To Measure Gait, Egress, and Locomotor Performance in Perturbed Environmental Conditions After Simulated Spaceflight Jeffrey Willey, Ph.D., Wake Forest School of Medicine, Winston-Salem, North Carolina



Spatiotemporal Mapping of the Impact of Spaceflight on the Heart and Brain Christopher Mason, Ph.D. Weill Medical College of Cornell University, New York, NY



Using Water Bears to Identify Biological Countermeasures to Stress During Multigenerational Spaceflight Thomas Boothby, Ph.D., University of Wyoming Laramie, Wyoming



The Effects of Spaceflight and Reloading on Skeletal Muscle and Bone Mary Bouxsein, Ph.D., Harvard Medical School, Boston, Massachusetts



Space Environment and Epigenetics of Endocrine Regulation of DNA Repair and Cell Cycle in Mammary Epithelial Cells

Donato Romagnolo, Ph.D., University of Arizona, Tucson, Arizona

### Featured Animal Biology Flight-based Projects



Megakaryocytes Orbiting in Outer Space and Near Earth: The MOON Study Hansjorg Schwertz, Ph.D. University of Utah, Salt Lake City, UT



Integrated physiological responses of CNS and muscle in Drosophila and C. elegans along a gravity continuum Karen Ocorr, Ph.D. Sanford Burnham Prebys. Medical Discovery Institute. La Jolla, CA



Effects of Spaceflight on Gastrointestinal Microbiota in Mice: Mechanisms and Impact on Multi-System Physiology Fred Turek, Ph.D., Northwestern University, Evanston, Illinois



UMAMI: Impact of Spaceflight on Beneficial Animal-Microbe Interactions Jamie Foster, Ph.D., University of Florida, Gainesville, Florida



Role of Mesenchymal Stem Cells in Microgravity Induced Bone Loss Abba Zubair, Ph.D., Mayo Clinic, Jacksonville, Florida

## Featured Plant Biology Spaceflight Projects



APEX-08: Can Polyamines Mitigate Plant Stress Response under Microgravity Conditions? Patrick Masson, Ph.D., University of Wisconsin, Madison, Wisconsin



APEX-09: C4 Photosynthesis in Space (C4Space) Pubudu Handakumbura, Ph.D., Pacific Northwest National Laboratory, Richland, Washington



APEX-11: Hypobaric Plant Biology in Space Exploration -Molecular Responses of *Arabidopsis* to Combined Effects of Low Atmospheric Pressures and Microgravity of Spaceflight Vehicles

Anna-Lisa Paul, Ph.D., University of Florida, Gainesville, Florida



BRIC-24: Membrane Contacts in Plant Gravity Perception Marcella Rojas-Pierce, Ph.D., North Carolina State University, Raleigh, North Carolina



 BRIC-27: From Antarctica to Space: Molecular Response and Physiological Adaptation of Moss to Simulated Deep Space Cosmic Ionizing Radiation and Spaceflight Microgravity.
 INSTITUTE Agata Zupanska, Ph.D. SETI Institute, Mountain View, California



BRIC-LED-002: BRIC: Exploring Spaceflight-Linked Changes in Plant Defense Capabilities Simon Gilroy, Ph.D., University of Wisconsin, Madison, Wisconsin







MVP-Plant-01: RNA Plant Regulation Redux (in MVP) Imara Perera, Ph.D., North Carolina State University, Raleigh, North Carolina

PH-03: Epigenetic Adaptation to the Spaceflight Environment - Accumulated Genomic Change Induced by Generations in Space

Anna-Lisa Paul, Ph.D., University of Florida, Gainesville, Florida

PH-08: Evaluation of Small Plants for Agriculture in Confined Environments (SPACE) Tomatoes for Space Flight Applications Robert Jinkerson, Ph.D., University of California, Riverside,

Robert Jinkerson, Ph.D., University of California, Riverside, California



Developing A System for Rapid Diagnosis of Plant Diseases and Monitoring of Plant Microbiome for Spaceflight Applications

Natasha Haveman, Ph.D. University of Florida, Gainesville, FL

## Featured Plant Biology Ground-based Projects



Suborbital: The Role of Ca2+ Signaling During the Early Events of Plant Adaptation to Spaceflight Rob Ferl, Ph.D., University of Florida, Gainesville, Florida



Antarctic EDEN ISS: Spectral Imaging within the EDEN ISS Project – An Antarctic Analog for Enhancing Exploration Life Support Rob Ferl, Ph.D., University of Florida, Gainesville, Florida



Leveraging Spaceflight Genomic Data to Uncover Developmental and Cell Type Specific Gene Regulatory Networks in Plants Responding to Gravity Simon Gilroy, Ph.D., University of Wisconsin, Madison, Wisconsin



RNA-Seq Guided Mutant Analysis to Discover New Components of Gravity Signaling in Plants Scot Wolverton, Ph.D., Ohio Wesleyan University, Delaware, Ohio



Modeling Leafy Greens Physiological and Biochemical Responses to Light Intensity and Successive Harvest Kellie Walters, Ph.D., University of Tennessee, Knoxville, Tennessee



Using Brachypodium distachyon to Study the Molecular Mechanisms that Underlie Directional Root-Growth Responses to Low-Speed 2-Dimensional Clinorotation Patrick Masson, Ph.D., University Of Wisconsin, Madison, Wisconsin



Time and Space, Determining the Interactions Between the Circadian Clock and Microgravity Colleen Doherty, Ph.D., North Carolina State University, Raleigh, North Carolina



The Use of Microgravity Simulators for a Mechanistic Understanding of Cytoskeletal-mediated Regulation of Root Growth

Simon Gilroy, Ph.D., University of Wisconsin-Madison, Wisconsin

# Extended Longevity of 3D Tissues and Microphysiological Systems for Modeling of Acute and Chronic Exposures to Stressors (POC: Lisa Carnell)

(Sponsored by NASA's Space Biology Program in partnership with NASA's Science Mission Directorate, NASA's Human Research Program (HRP), NIH's National Center for Advancing Translational Sciences (NCATS), the NIH National Institute of Allergy and Infectious Diseases (NIAID), the NIH National Cancer Institute (NCI), BARDA, and the FDA).



Understanding the Brain-Liver-Gut Axis during Spaceflight and Aging. Elizabeth Blaber, Ph.D, Rensselaer Polytechnic Institute



Identification of Biomarkers and Pathological Mechanisms via Longitudinal Analysis of Neurological and Cerebrovascular Responses to Neurotoxic Stress Using a Multi-cellular Integrated Model of the Human Brain

Joel Blanchard, Ph.D. ICAHN School of Medicine at Mount Sinai



Bioengineer Long-lasting 3D Neurovascular Microphysiological System to Model Chronic Inflammation-mediated Neurodegeneration Guohao Dai, Ph.D. Northeastern University



Long-term Patient iPSC Vessel Chip Model to Assess Stressors of Atherosclerosis and mRNA Therapeutics Abhishek Jain, Ph.D. Texas A&M



Long-lived Single- and Multi-organ Tissue Equivalent (OTE) Platforms to Model the Response of Human Tissues to Various Stressors Christopher Porada, Ph.D. Wake Forest University



MORPH: Multi-Organ Repair Post Hypoxia Gordana Vunjak-Novakovic, Ph.D. Columbia University



Assessing Long-Term Effects of Radiation Exposure in Engineered Heart & Vascular Tissues Joseph Wu, MD. Ph.D. Stanford University



Extended Culture of Kidney MPS and Organoids to Model Acute and Chronic Exposure to Drugs and Environmental Toxins Catherine Yeung, Pharm D. Ph.D. University of Washington

## KSC ISS Payload Manifest – Increments 67-70

Increment Flight					67/68	69/70							
		Crew-4	SpX-25	Crew-5	NG-18	<b>SpX-26</b> (possibly 45- day mission)	SpX-27	NG-19	USCV-6	<b>SpX-28</b> (possibly 45-day mission)	USCV-7	NG-20	SpX-29
Launch		4/27/22	7/15/22	10/5/22	11/7/22	11/21/22	2/19/23	3/11/23	2/19/23	6/1/23	8/17/23	Oct 2023	Dec 2023
Unc	lock	10/14/22	8/19/22	2/28/23	1/27/23	1/7/23	3/22/23	7/1/23	8/27/23	7/18/23	Feb 2024	Jan 2024	Jan 2024
6	Ascent	-	DynaMoS (Jansson)		PH-03A (Paul)	Veg-05 (Massa) BRIC-26 (Nicholson)	APEX-10 (Gilroy)			APEX-09 (Handakumbura) BRIC-25 (Rice) BRIC-LED-002 (Gilroy) PH-03B (Paul)			Veg-06 (Lewis)
Payloads	Descent	XROOTS (plant samples)	MVP Plant01 (Run 2 modules)			DynaMoS (Jansson) BRIC-26 (Nicholson) XROOTS (plant samples; 4th harvest)	APEX-10 (Gilroy) PH-03A (Paul) XROOTS (hardware)			APEX-09 (Handakumbura) BRIC-25 (Rice) BRIC-LED-002 (Gilroy) Veg-05 (Massa)			PH-03B (Paul)

## ARC ISS Payload Manifest – Increments 67-70

Increment Flight Launch		67/68								69/70					
		Crew-4	<b>SpX-25</b> 7/15/22	Crew-5	NG-18	<b>SpX-26</b> (possibly 45- day mission) 11/21/22	<b>SpX-27</b> 2/19/23	NG-19 3/11/23	USCV-6 2/19/23	SpX-28 (possibly 45-day mission) 6/1/23	USCV-7 8/17/23	NG-20 Oct 2023	SpX-29		
		4/27/22		10/5/22	11/7/22								Dec 2023		
Und	ock	10/14/22	2 8/19/22	2/28/23	1/27/23	1/7/23	3/22/23	7/1/23	8/27/23	7/18/23	Feb 2024	Jan 2024	Jan 2024		
							MHU-8* (HRP/SB)			MoSL-2 (O'Connell)		EnteroGAIT (Carr)	RR-26* (Ronca)		
	Ascent									MVP-Cell-02A** (Everroad)					
S	A	-								RR-20* (Christenson)					
Payloads			RR-18 (Mao)				MHU-8* (HRP/SB)			MoSL-2 (O'Connell)		EnteroGAIT (Carr)	RR-26* (Ronca)		
	Descent									MVP-Cell-02A** (Everroad) RR-20*					
										(Christenson)					

# EXPLORE MOONtoMARS

NASA

### Interplanetary space radiation What are we going to encounter beyond Low Earth Orbit (LEO)?



Galactic Cosmic Rays

**Trapped** Particles

Solar Protons

Heavier Ions

Ionizing radiation that will affect biology:

- Galactic Cosmic Radiation (GCR)
- Solar Particle Events (SPEs)

Limits of life in space, as studied to date:

- ♦ 12.5 days on a lunar round trip
- ♦ 1.5 years in low Earth orbit on ISS

### Radiation Environment of the Moon: Lunar surface radiation is a distinct environment from deep space



Image credits: NASA

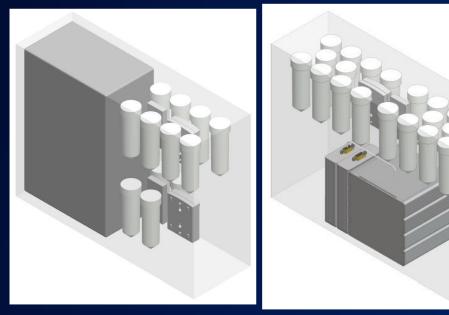
Lunar regolith absorbs galactic cosmic radiation (GCR) to produce secondary albedo neutrons

- High-energy, fast neutrons cause direct cellular damage
- Fast neutrons also produce ionizing radiation
- Estimates of surface neutron doses vary widely

## BioExpt-1: Flying on Artemis-1

Anticipated Launch: November 2022

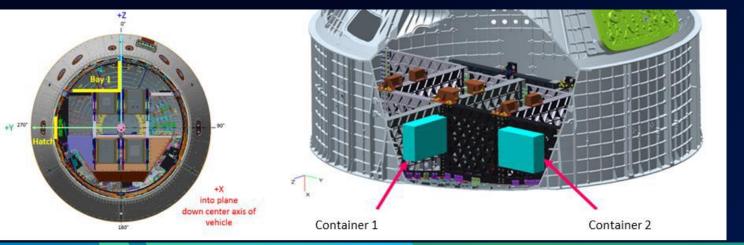
- Plant seeds Arabidopsis thaliana: Life Beyond Earth: Effect of Space Flight on Seeds with Improved Nutritional Value; PI Federica Brandizzi, Michigan State University
- Algae Chlamydomonas reinhardtii: Fuel to Mars; PI Timothy Hammond, Institute for Medical Research; Chlamydomonas reinhardtii
- Fungi Aspergillus niger: Investigating the Roles of Melanin and DNA Repair on Adaptation and Survivability of Fungi in Deep Space; PI Zheng Wang, Naval Research Laboratory
- Yeast Saccharomyces cerevisiae: Multi-Generational Genome-Wide Yeast Fitness Profiling Beyond and Below Earth's Van Allen Belts; PI Luis Zea, University of Colorado-Boulder



SBP Payload Container Assembly configuration for Drs. Zea & Brandizzi (left) and Drs. Wang & Hammond (right).



3D printed ½ scale mockup of SBP Payload Container Assembly with blue Science Bag contained within.



The SBP Payload Container Assemblies are installed onto backbone panels 1 and 2 in Bay 1 of the Orion capsule.

## Lunar Explorer Instrument for Space Biology Applications (LEIA)

- LEIA is based on the 6U BioSentinel ۲ Small Sat
- Modified 4U Biosensor with a 2U • "Pseudo Bus" to provide thermal control for the lunar surface and data conditioning





Accommodates yeast and related cellular systems

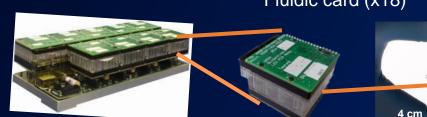
### **4U BioSensor Payload**

(2U Pseudo Bus not shown in diagrams below)

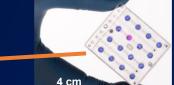


LET spectrometer attached at one end of payload





Fluidic card (x18)



Biosensor payload with fluidic cards and optical sensors

## LEIA: Lunar Explorer Instrument for space biology Applications A Funding opportunity released as Program Element E.10 in ROSES 2021 on NSPIRES

(Selections Announced: September 3, 2021)

We solicited proposals for an autonomous biological research experiment using yeast as a model organism as part of the project-formulation phase in the work of the Lunar Explorer Instrument for space biology Applications (LEIA) call



- Solicited for experiments to be integrated into the Biosentinel hardware and be tested on the • ground to establish a functional experimental payload that could be flight-ready in future.
- The overarching goal of the proposed research studies themselves were to investigate the • effects of lunar environmental conditions on yeast biology, however, proposals are to be written with the understanding that the LEIA project is still in its ground-based testing

#### Three investigations were selected:



Investigating Lunar Stress and Parkinson's Disease using an Alpha Synuclein Yeast Model Lynn Harrison, Ph.D. Louisiana State University System,

Shreveport, LA





**ORGANA: Oxidation-Reduction potential and Genetic Assessments** for New mission Applications Sergio Santa Maria, Ph.D. NASA Ames Research Center, Mountain View, CA

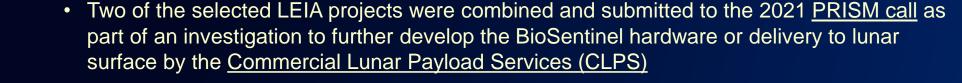


Feasibility of synthetic biology countermeasures for human exploration beyond low Earth orbit Andrew Settles, Ph.D. NASA Ames Research Center, Mountain View, CA

## PRISM: Payloads and Research Investigations on the Surface of the Moon

This Funding opportunity was released by NASA ESSIO (Exploration Science Strategy & Integration Office) (Selected July 2, 2022)

Investigations that include development and flight of science-driven suites of instruments payloads that will be delivered to the lunar surface by the Commercial Lunar Payload Services (CLPS)





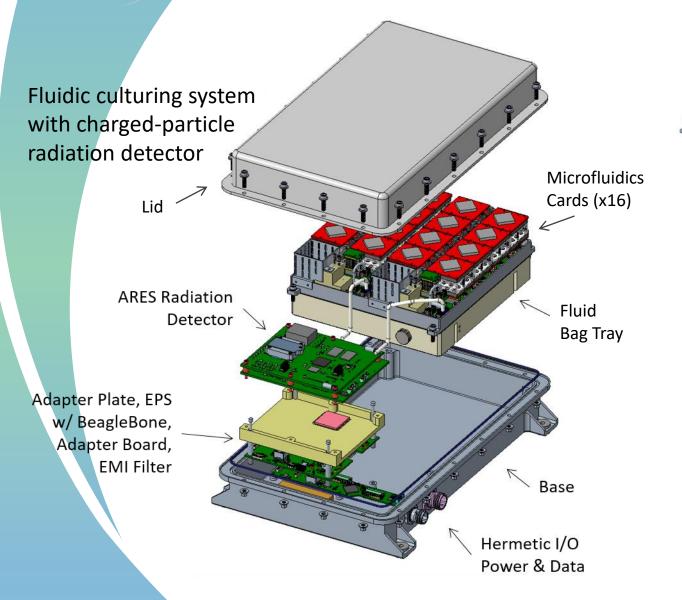
ORGANA: Oxidation-Reduction potential and Genetic Assessments for New mission Applications Sergio Santa Maria, Ph.D. NASA Ames Research Center, Mountain View, CA

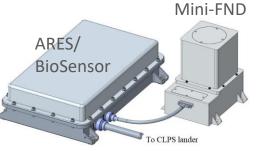
Feasibility of synthetic biology countermeasures for human exploration beyond low Earth orbit Andrew Settles, Ph.D. NASA Ames Research Center, Mountain View, CA

 Both investigators and the Biosentinel hardware were selected for a CLPS flight as part of a future Artemis Mission



# The LEIA-PRISM Experiment

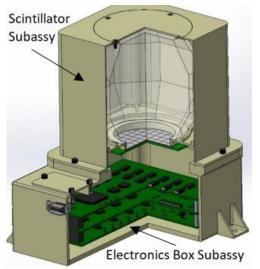




\*MLI not shown

## Mini-FND

fast neutron detector



#### Approach:

- Engineer yeast strains to test growth and metabolism for sensitivity to radiation.
- Measure biologically relevant radiation in transit and on the lunar surface.

#### Expected Outcomes/Impact:

- Determine cellular sensitivity to the lunar environment.
- Evaluate feasibility of bioproduction on the Moon.
- Test genetic strategies to enhance cellular tolerance to the lunar environment.
- Determine radiation risks for crew.

## Science Working Group

#### • BLISS (Beyond LEO Instrumentation and Science Series)

- Consists of members of research community representing subject matter experts in fundamental Space Biology focusing on diverse model organisms
- Skewed heavily external from research institutions, with support from NASA internal SMEs mostly in facilitator roles
- Meets monthly to discuss future research needed to be conducted BLEO, and what capabilities are required to support

#### • 2021 Annual Report released 10/15/21

- Covers opinion of SWG community on what research can and should be conducted in BLEO environment in next 5 years
- <u>https://ntrs.nasa.gov/citations/20210023324</u>

#### • 2022 Annual Report to be released mid-November

- Expands the focus 5-15 years into future
- Incorporates AI/ML to enable mission execution and enhance science return

#### • Lunar Biology Technology (LBTech) Workshop - April 20-21, 2022

- Facilitated discussion between Science community, Payload Developers, and Platform Providers to strategize on BLEO research and infrastructure
- LBTech 2 coming next Spring (2023)

National Aeronautics and Space Administration

Space Biology Beyond LEO Instrumentation & Science Series Science Working Group 2021 Annual Report

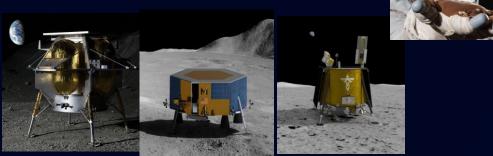
R. Craig Everroad<sup>1</sup>, Jamie Foster<sup>2</sup>, Jonathan M. Galazka<sup>1</sup>, Janet Jansson<sup>3</sup>, Jessica A. Lee<sup>1</sup>, Matthew P. Lera<sup>1</sup>, Imara Perera<sup>4</sup>, Antonio Ricco<sup>1,5</sup>, Nathaniel Szewczyk<sup>6</sup>, Paul Todd<sup>7</sup>, Ye Zhang<sup>8</sup>, Lynn Harrison<sup>9</sup>

NASA Ames Research Center, 2. University of Florida, 3. Pacific Northwest National Laboratory,
 North Carolina State University, 5. Stanford University, 6. Ohio University, 7. TechShot, Inc.,
 NASA Kennedy Space Center, 9. Louisiana State University Health Shreveport

# Future Needs for Science Beyond LEO

# Capabilities for Animal & Model Organism Studies on the

## Moon



Taking science to the lunar surface





#### Habitats & analytical tools for research

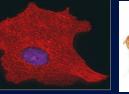




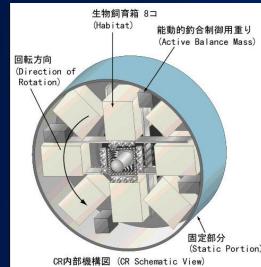
Use well-characterized animal model organisms & cellular systems to understand the complex biological consequences of exposure to deep space's unique environment.

- Facilities/equipment/habitats required for lunar surface experimentation with animals and related systems, need to be developed for Space Biology.
- In-flight entrifuges that can simulate 1g or fractional g loads, can be used to differentiate between the biological effects of altered gravity vs other stressors like radiation.





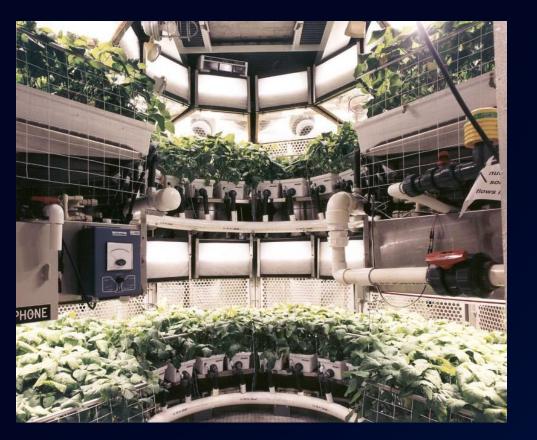




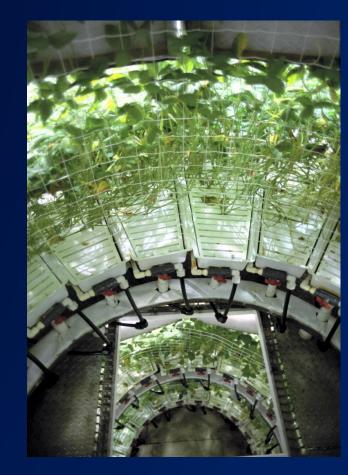




## Growing Plants on the Moon



Develop a plant growth facility on the lunar surface to understand the complex biological consequences of exposure to deep space's unique environment and validate the ability to produce supplemental nutrients for future space explorers



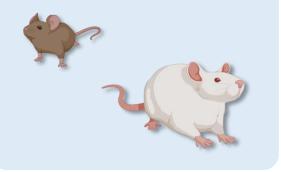
# Accomplishments in Open Science

## **Open Science Projects**

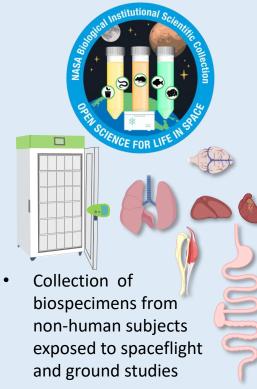
Biospecimen Sharing Program (BSP)



- Dissects and preserves rodent tissues from Flight and Ground investigations
- Coordinates internal tissue sharing



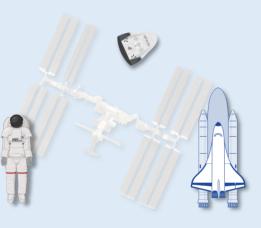
NASA Biological Institutional Scientific Collection (NBISC)



 Coordinates and fulfills biospecimen requests and awards Ames Life Sciences Data Archive (ALSDA)



 Collects and curates physiological, mission, and imaging data



GeneLab (GL)



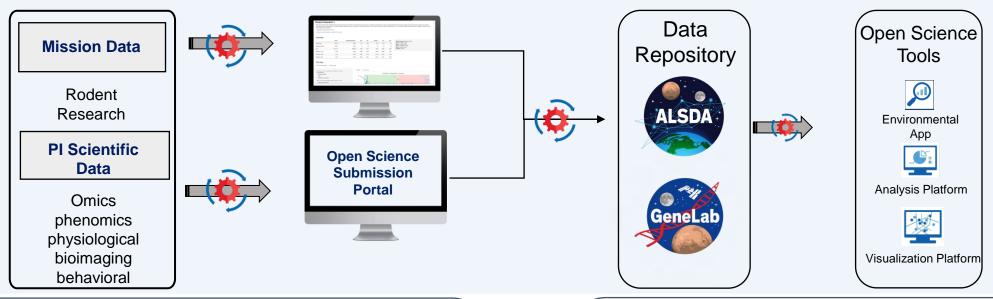
 Collects and curates omics data from space-relevant samples



## **Biological Data Repository**

### ALSDA and GeneLab Data Systems Integration (2022)

The Open Science Data Repositories will introduce a multi-project web-based submission portal to support self-service metadata curation and data submission within FAIR Guidelines.



### AI/ML 4 Life in Space (SMD Funded)

- Ensure data is AI/ML ready
- Generate benchmark datasets to train new AI/ML algorithms
- Utilize current AI/ML tools to analyze space biology data

### **Analysis Working Group (AWG)**

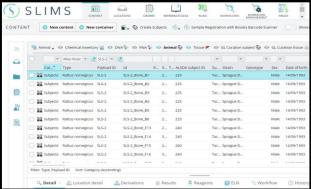
- 400+ members from multiple space agencies, international institutions, and industry
- Scientists meet monthly with each group to provide feedback, standards and analyze data
- Five Groups: Animal, Microbes, Plants, Multi-Omics, ALSDA
- Published 11 collaborative peer-reviewed papers to date

## Biological Specimen Sharing Program (BSP) FY22 Accomplishments

- Rodent Research-12 (RR-12) BSP
  - Dissections successfully completed at Loma Linda University (California) October 11-26, 2021, after postponement from March 2020 due to COVID-19.
  - ~3,500 biospecimens were collected and transferred to NBISC.
- Rodent Research-18 (RR-18) BSP
  - Live Animal Return dissections successfully completed at The Roskamp Institute (Florida) in January 19 February 7, 2022, after SpX-24 return.
  - Readaptation group dissections successfully completed at Loma Linda University April 16 May 9, 2022.
  - ~12 tissues types are available in NBISC for the greater research community to request.
- Ground investigation: Bone Remodeling Under Differential Gravitational Environments (Komatsu) BSP
  - Remote rat dissections (3 of 8) successfully completed at Stony Brook University (New York) on May 5, May 31, and August 15.
  - 5 tissues types will be available in NBISC for the greater research community to request.
- Space Radiation Element/Space Biology Collaboration
  - Pilot study initiated between Space Radiation Element, BSP, and NBISC to ingest biospecimens and data from SR PI Dr. Honglu Wu into NBISC.
  - Over 7 tissue types will be available in NBISC for the greater research community to request. Data entry into the Laboratory Information Management System completed in August 2022.
  - Joint BSP/NBISC/ALSDA/GL presentation completed at the inaugural Space Radiation Biospecimen & Tissue Sharing Summit on September 13, 2022, where over 140 participants engaged in discussions about open science and specimen and tissue sharing.
- BSP and NBISC Biospecimen Data Migration to Cloud
  - ~32,000 biospecimen records transferred into the shared, cloud-based Laboratory Information Management System to automate workflows and better manage sample metadata. Completed in September 2022.

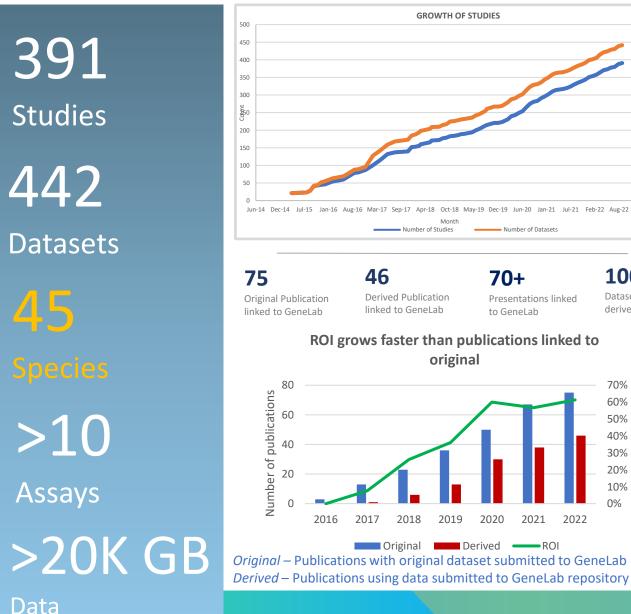






## **GeneLab Metrics**

In FY22, GeneLab has added 56 studies publicly for access worldwide.



#### New features on the Data Systems

- The visualization portal has now expanded to include Gene Set Enrichment Analysis (GSEA) plots. GSEA is an analysis method that determines whether a defined set of genes shows statistically significant and concordant differences between two biological states.
- GeneLab Repository gets a new user interface with a ٠ new design and additional search capabilities to enable discovery.

#### Science Communication & Outreach

100 +

60%

30% 20%

10%

0%

Datasets used in

Origina 40%

Derived/

derived publications

On June 30<sup>th</sup>, GeneLab hosted the 2022 AWG Symposium from 10 am – 2:30 pm PT. This virtual event allowed the pubic an opportunity to view some exciting omics science stemming from the AWG community and was attended by 350 unique participants. During the event, attendees had an opportunity to sign-up for the AWGs which led to 45 new members, including early career scientists and university professors, resulting in over 400 AWG members to date.

## GeneLab Expands Bioinformatics Training to HBCUs and MSIs

 GeneLab for Colleges and Universities (GL4U) is partnering with Jet Propulsion Laboratory's (JPL) Planetary Protection Center of Excellence to provide space biology-relevant training in bioinformatics to educators at historically black colleges and universities (HBCUs) and minority serving institutions (MSIs).

 GL4U: RNAseq Educator Bootcamp was held virtually from May 31<sup>st</sup> - June 10<sup>th</sup>, 2022, for 6 educators and 4 graduate students.

- One student, Chiefe Mo, was so inspired that he composed and performed a GeneLab anthem on the last day - https://www.youtube.com/watch?v=szC3y2HWFcQ
- During the bootcamp, educators received training via lectures and hands-on instruction using Jupyter Notebooks (JNs) on how to 1) analyze GL RNAseq data and 2) run the RNAseq bootcamp for students at their home institutions, thereby extending the reach of this initiative to more students.
- To ensure educators from underserved colleges and universities are able to successfully implement the RNAseq training at their home institutions, the required computer resources will also be made available to them through the NASA Science Managed Cloud Environment (SMCE) funded by SMD AWS Space Act Agreement.





Dr. Tyesha Farmer Dr. Elba Serrano Assistant Professor of Genetics Regents Professor of Biology Assistant Professor of Chemistry Alabama A&M University New Mexica State University

Dr. Keneshia Johnson Alabama A&M University







Dr. Rachel Mackelprang Professor of Microbiology California State University (CSU), Northridge

Dr. Wei-Jen Lin Professor in the Dept. of **Biological Sciences** California State Polytechnic University

Assistant Professor in the Dept. of Biological Sciences California State Polytechnic University









Dr. Joel Steele Chiefe Mo **Fulbright Future** Scholar, Proteomics California State Monash University

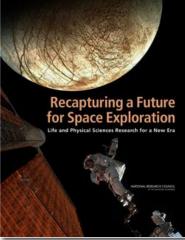
Master's Student, Biology Biology Polytechnic University

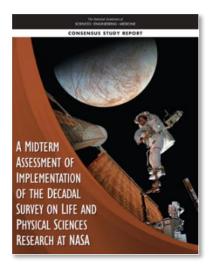
Suzi Arzoumanyan Mark Ortiz Teaching Associate. Master's Student, Biology California State CSU, Northridge Polytechnic University

# Recent and Future Space Biology Funding Opportunities

# 2023-2032 Decadal Survey for Biological and Physical Sciences Research in Space

- A National Academies of Sciences, Engineering, and Medicine report
  - Purpose is to generate consensus recommendations for a comprehensive vision and strategy for a decade of transformative science
  - Focused, select number of highest priority recommendations
  - Expected to be delivered in the summer of 2023
- NASA's Decadal response plan
  - Share initial plan at a Town Hall ~90 days post receipt
  - Formulate a phased approach to implementation
    - Identify initial activities starting in FY24
    - Propose Decadal response budget beginning FY25
    - Staggered start of flight programs
- Current activities while awaiting new Decadal
  - Identifying commercial space capabilities and science community needs to accelerate the pace of research in space
  - New solicitations in focus areas (Quantum Science, Thriving in Deep Space (TIDES))





### 2022's Space Biology Annual Ground/Flight Funding Opportunities

Space Biology Plant Studies Released as <u>Program Element E.9 in ROSES 2021</u>: Released: December 16, 2021; Final Proposals due: April 28, 2022 Selections Announced: Late October 2022

- Released in collaboration with the NASA Human Research Program (HRP)
  - Space Biology will make and fund awards to investigators through E.9, while the Space Radiation Element in HRP will provide funded investigators access to the NASA Space Radiation Laboratory (NSRL) at Brookhaven National Laboratory for ground radiation studies
- Science Emphases:
  - Advance the understanding of how plants accommodate to the spaceflight environment (e.g., alterations in gravity, radiation, elevated carbon dioxide levels, etc.).
  - Study spaceflight environmental effects on plant physiology, biochemistry, cellular, and molecular biology during development, growth, and/or propagation.
  - Understanding biological processes that will enable NASA's long-term goal of sustained agriculture
  - HRP is interested in the psychological and nutritional benefits for astronauts of growing plants to support long duration human exploration to deep space destinations such as Mars.

Awards were made to seven investigators from seven institutions in six states. Total value of awards = ~\$1.7M

### 2022's Space Biology Annual Ground/Flight Funding Opportunities (Cont)

Space Biology Animal Studies Released as <u>Program Element E.11 in ROSES 2021</u>: Released: December 16, 2021; Final Proposals due: April 21, 2022 Selections Announced: Late October 2022

- This program element was released in collaboration with the NASA Human Research Program (HRP)
- Science Emphases:
  - Fundamental questions that will advance the understanding of how animal systems accommodate to the spaceflight environment for important stressors such as ionizing radiation and/or changes in gravity.
  - Study spaceflight environmental effects on animal physiology, biochemistry, cellular, and molecular biology responses.
  - Emphasis on physiological systems that are known to respond to the spaceflight and radiation environments (e.g., neurobehavioral, cardiovascular, immune, musculoskeletal, etc.) are encouraged especially those with relevance or translation to the human system.
  - The use of omics/systems biology type approaches that examine the effects of radiation, altered gravity, or other stressors encountered in the spaceflight environment are encouraged.
  - Understanding biological processes that will enable NASA's goal of long-duration human exploration to deep space destinations such as Mars.

Awards were made to twelve investigators from eight institutions in seven states. Total value of awards = ~\$3.5M

### Research Pathfinder for Beyond Low Earth Orbit Space Biology Investigation (Artemis II) Funding Opportunity

#### Released as **Program Element E.11 in ROSES 2022**:

Released: August 2022; Final Proposals due October 13 (+2 weeks extension due to hurricane), 2022 Anticipated Selection Announcement: Early 2023

- This opportunity solicited proposals for biological research experiments with invertebrates to be conducted on the Artemis II
  mission.
- The goal of these projects will be to study early changes in physiological systems due to exposure to the deep space environment.
- Selection of projects for this opportunity will be contingent on whether the Artemis II mission is able to accommodate the
  program's biological payloads on the Orion spacecraft

NASA anticipates selecting two projects for award, each with a maximum budget of \$750K

### 2021 Collaborative 3D Tissue Model Funding Opportunity

### Extended Longevity of 3D Tissues and Microphysiological Systems for Modeling of Acute and Chronic Exposures to Stressors

Released: May 27, 2021: Final Proposal Due: September. 14, 2021 Selections Announced: March 1, 2022

### **POC: Lisa Carnell**

#### A collaboration between NASA's Space Biology Program and:

- NASA's Human Research Program;
- National Institutes of Health's National Center for Advancing Translational Sciences (NCATS);
- National Cancer Institute (NCI);
- National Institute of Allergy and Infectious Diseases (NIAID);
- Biomedical Advanced Research and Development Authority (BARDA);
- Food and Drug Administration (FDA);

Purpose of studies are to extend the current longevity of existing "tissue chip" or "organs-on-chip" systems, which will have utility for understanding human disease pathology, acute, chronic and repeated drug or toxin exposures, and exposures to environmental hazards, both on Earth and in space.

• Studies are for a maximum 4-years with a maximum budget of \$2M over the duration of the study

Awards were made as contracts to eight investigators from eight institutions in six states. Total value of awards = ~\$16M

### **Space Biology's Upcoming ROSES Solicitations:**

### FY2022 Annual Ground and Flight Research Announcement

To be Released under ROSES-2022 Release: ~Late Fall / Winter 2022, Proposals Due: ~Spring 2021 Selections Announced: ~Late Summer/Early Fall 2023

Solicited Topics: TBD

# Challenges

# Thoughts:

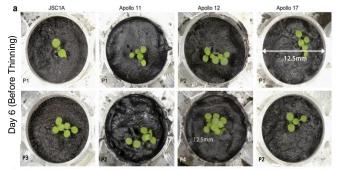
- 1. Total available \$ to award Space Bio grants have reduced in the past 2 years
- 2. We do need to continue the capability to do Space Biology research in low Earth orbit even after the ISS is phased out since it will be a while before we can conduct complex experiments (e.g., rodent research etc.) beyond LEO.
- 3. Ground research will continue to be important alongside flight research as we test the effects of multiple deep-space stressors on different platforms on the ground as well as in flight. Therefore, the requisite facilities need to be maintained.
- 4. It will be important to develop the capabilities to conduct long duration experiments in the lunar environment in preparation for Mars missions: need to do comparative science between different spaceflight platforms and between different organisms.
- 5. We need to grow as a program to be able to start taking full advantage of deep-space exploration opportunities.
- 6. There is no decadal wedge funding set aside for us to respond to the upcoming recommendations.

# Recent Science Highlight

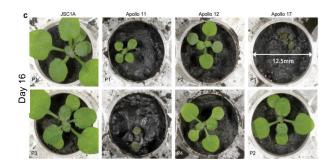
Plants grown in Apollo lunar regolith present stress-associated transcriptomes that inform prospects for lunar exploration. <u>Paul AL</u>, Elardo SM, <u>Ferl R</u>. Commun Biol. 2022 May 12;5(1):382. DOI: <u>10.1038/s42003-022-03334-8</u>

### Arabidopsis thaliana germinates and grows in regolith

- Regolith samples from Apollo 11, 12, and 17 were used.
- JSC-1A = simulant control.

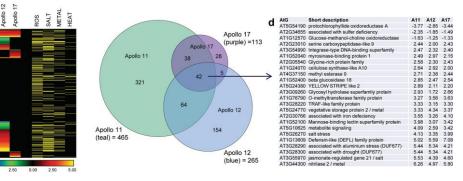


• Germination rates were close to 100% in all sources of Apollo regolith.



• Regolith-grown seedlings, however, did not thrive compared to JSC-1A controls.

### Plant transcriptomes differ by Apollo samples (site)



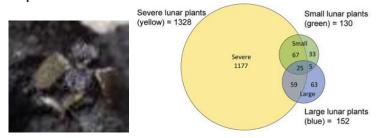
Expression data were parsed based on lunar sample site replicates compared to controls

### Plant transcriptomes differ by morphology

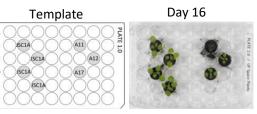
 Plants with a "severe" phenotype (tiny, abnormal morphology and reddish-black pigmentation) had 1000+ DEGs, demonstrating severe reactions to the regolith. Anna-Lisa Paul Robert Ferl

Univ. of Florida, Gainesville

- All regolith samples, irrespective of Apollo site, significantly evoked differentially expressed genes (DEGs) indicative of a strong stress response.
- 71% of the DEGs typically were associated with salt, metal, and reactive oxygen species stress.



**Relevance:** Terrestrial plants can grow in lunar regolith, however, further characterization and optimization is required before regolith can be considered a routine in situ resource, as it is not a benign growth substrate.



# Conclusion

### • As NASA plans to...

- return to the lunar surface,
- develop sustainable lunar habitation, and
- prepare to explore Mars

### • Space Biology intends to...

- utilize multiple biological model systems and
- spaceflight platforms

> To understand the mechanisms of change in biological systems in response to long duration exposure to deep space,

➤To enable exploration,

> To benefit life on Earth: human health & controlled environment agriculture.



# **Backup Slides**

# **Upcoming BSP Objectives**

- Rodent Research-18 (RR-18) BSP Completion
  - Frozen carcass dissections at Loma Linda University (California), October 23-November 1, 2022.
  - Dr. Mao: Space Flight Environment Induces Remodeling of Vascular Network and Glia-vascular Communication in Mouse Retina.
- NASA JAXA Joint Partial-gravity Rodent Research (JPG-RR), Mouse Habitat Unit-8 (MHU-8)
  - Fresh dissections at KSC NET February 2023 and March 2023.
  - Dr. Bouxsein (HRP), Dr. Fuller (SB), Dr. Vitaterna (SB), Dr. Takahashi (JAXA).
- Rodent Research-20 (RR-20)
  - Fresh carcass dissections at The Roskamp Institute (Florida) NET March 2023 and frozen carcass dissections at University of Kansas Medical Center NET July 2023.
  - Dr. Christenson: Female Reproductive Health: Space Flight Induced Ovarian and Estrogen Signaling Disfunction, Adaptation, and Recovery.
- Rodent Research-26 (RR-26) BSP
  - Fresh and frozen carcass dissections at KSC NET July 2023.
  - Dr. Ronca: Rodent Research Habitat Validation 2.
- Komatsu Ground Study BSP Completion
  - Fresh carcass dissections at Stony Brook University (New York) October-July 2023.
  - Dr. Komatsu: Bone Remodeling Under Differential Gravitational Environments.
- Data Collection Optimization between BSP, NBISC, and GeneLab
  - Implementation of an automated vial barcoding system and hardware to expediate labeling and processing of samples.
  - Integration across SB Open Science will streamline interoperability and end-user analysis.

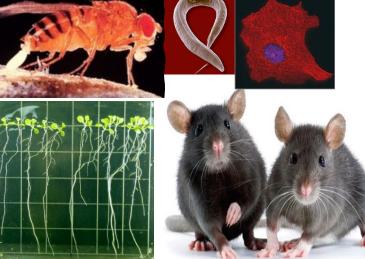












## **Combined Effects of Spaceflight Stressors**

#### Spaceflight stressors can include:

- Elevated levels of ionizing radiation
- Altered gravity conditions
  - Microgravity, hypergravity, partial gravity on planetary surface
- Elevated CO2 levels; altered PO2
- Exposure to regoliths/dust
- Altered day/night light cycles
- Social isolation, etc.

lonizing radiation and altered gravity are two important, spaceflight stressors that show various responses when applied together, depending on the organism, tissue type and assay. When combined, a given response to stressor may be:

- Amplified (additive, synergistic)
- No effect
- Suppressed (data not shown)

### **Bottom line:**

We still have a *very limited understanding of how living organisms respond to the combinations of the various spaceflight stressors.* 

It is *critical to expand our knowledge base*, enabling us to better prepare for possible risks to astronaut health posed by long-duration spaceflight missions to the Moon and Mars.

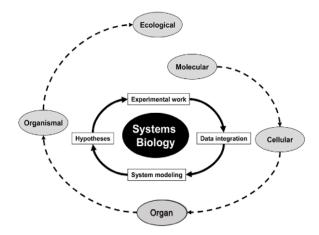
# <u>Thriving in Deep Space (TIDES)</u>



Transformative	1. Animal Biology	Vertebrate and invertebrate models to probe analogous changes in humans	
biological science and exploration applications	2. Plant Biology	From plant models to crops to sustain life for long-term human habitation	
	Microbiology	How it influences animals and plants in space	

### **Challenges of Interest for the future**





Systems Biology: Summing the component parts to create a model for predicting how the system will change over time and under varying environmental conditions. This approach could be applied to multiple biological systems to develop predictive models of the effect of long-duration spaceflight.

Effects of combined spaceflight stressors: e.g., deep-space radiation and altered gravity

#### **Research Campaign:** Are there research campaigns to consider?

Use well-characterized biological model organisms to understand the complex biological consequences of exposure to deep space's unique environment:

- Approach: Conduct a sequence of Earth-based, low Earth orbit (LEO), and lunar/cis-lunar investigations (in the next decade & more), with transit to and on Mars research as the long-term goal (in the 2030's timeframe). Populations of biological model organisms of increasing complexity will be used to gather data on changes in the underlying responses to space-unique environmental stressors, which do not occur in everyday Earth-based systems. The data from whole organisms, cellular, and organ systems will be compared within populations and across organisms to create a database that ultimately enables predicting physiological responses of humans to the deep space environment.
   Facilities/equipment/hardware/habitats required for lunar/cis-lunar spaceflight experimentation needs to be developed for Space Biology.
- **Resources for ground-based studies** could include ionizing radiation facilities (see slide 21), NASA's Space Radiation laboratory at Brookhaven National Laboratories (BNL), hypergravity facilities, and simulated microgravity platforms. Flight studies could compare results between ground (1g, baseline radiation), the ISS (microgravity, low radiation environment), autonomous free flyers (microgravity, high radiation environment), and lunar surface (1/6g, high radiation environment), in long-duration multigenerational studies with biological model organisms. Appropriate radiation sensors and flight hardware would be developed for free flyers and lunar missions.



### **Challenges of Interest for the future** (continued)

#### Research Campaign: Are there research campaigns to consider?

Develop a plant growth facility on the lunar surface to understand the complex biological consequences of exposure to deep space's unique environment and validate the ability to produce supplemental nutrients for future space explorers.

Approach: Develop a pressurized modular facility on the Moon to grow and test plants with automated health monitoring, sensor technologies, water recovery, nutrient and water delivery systems.

- Utilize genetic engineering, quantitative genomics, and other tools to test and optimize different plant types for suitability to the deep space environment.
- Study plant microbial interactions, stability and reliability
  of the plant growth systems at a larger scale than we currently have on the ISS.
- Assess the use of lunar regoliths as a growth medium.
- Evaluate techniques for biomanufacturing of pharmaceuticals, and other valuable bioproducts for long term human exploration.
- Resources for ground-based studies could include ionizing radiation facilities (see next slide), and NASA's Space Radiation laboratory at Brookhaven National Laboratories (BNL).
- Partnerships with internationals could further help with the infusion of new technologies, capabilities, and resources.

### Challenges of Interest for the future (continued)

# Supporting Research Capabilities: Are there large hardware facilities to consider?

#### Ground Facilities: Gravity Radiation PlanEt Simulator (GRAPES)

Ground facility that replicates conditions on the Moon and Mars to conduct realistic ground studies mimicking the type of environment that organisms (and supporting hardware) will have to endure under those harsh conditions. Large environmental chambers with chronic low dose-rate radiation capabilities, planetary conditions such as light cycle, simulated partial gravity, exposure to lunar/martian dust simulants, vibration, acoustic, circadian disruption, and altered atmospheric conditions (CO2, O2, humidity, temperature) as experimental cofactors.

#### Approach:

The implementation sites could include centrifuges and simulated microgravity capabilities to simulate gravity as a continuum combined with chronic gamma irradiation sources of low-LET low dose-rate ionizing radiation, and opportunities where similar gravity simulators can be used in conjunction with low dose-rate of simulated high-LET radiation. [Note: BNL is unable to provide chronic doses of low dose-rate high LET radiation].

#### Lunar Science Facilities:

Develop science capabilities for use on the Lunar surface like we currently have for the ISS: Multiorganismal facilities with habitats, automated science capabilities, conditioned environments, video cameras, lighting, radiation and other sensors etc. to test biological systems/organisms and microphysiological systems responses, in the absence or presence of radiation shielding, lunar dust exposure, 1g centrifugation, etc.



# ALSDA AWG Highlight

The ALSDA AWG contributed their expertise throughout the year to establish a consensus on standards for phenotypic experiments including assay, sample, and study design metadata which must be collected from PIs to ensure data reusability.

### ALSDA

Configurations

Assay Metadata

# Assay Configurations developed through AWG

- Behavior (Novel Object Recognition)
- Bone Microstructure (MicroCT)
- Behavior (Gait)
- Behavior (Elevated Plus Maze)
- Molecular/Cellular Imaging (Lt/Fluor Microscopy)
- Behavior (Open Field)
- Immunoassay
- Flow Cytometry (Flow Cytometry)
- Calcium Uptake
- And more upcoming . . .

