

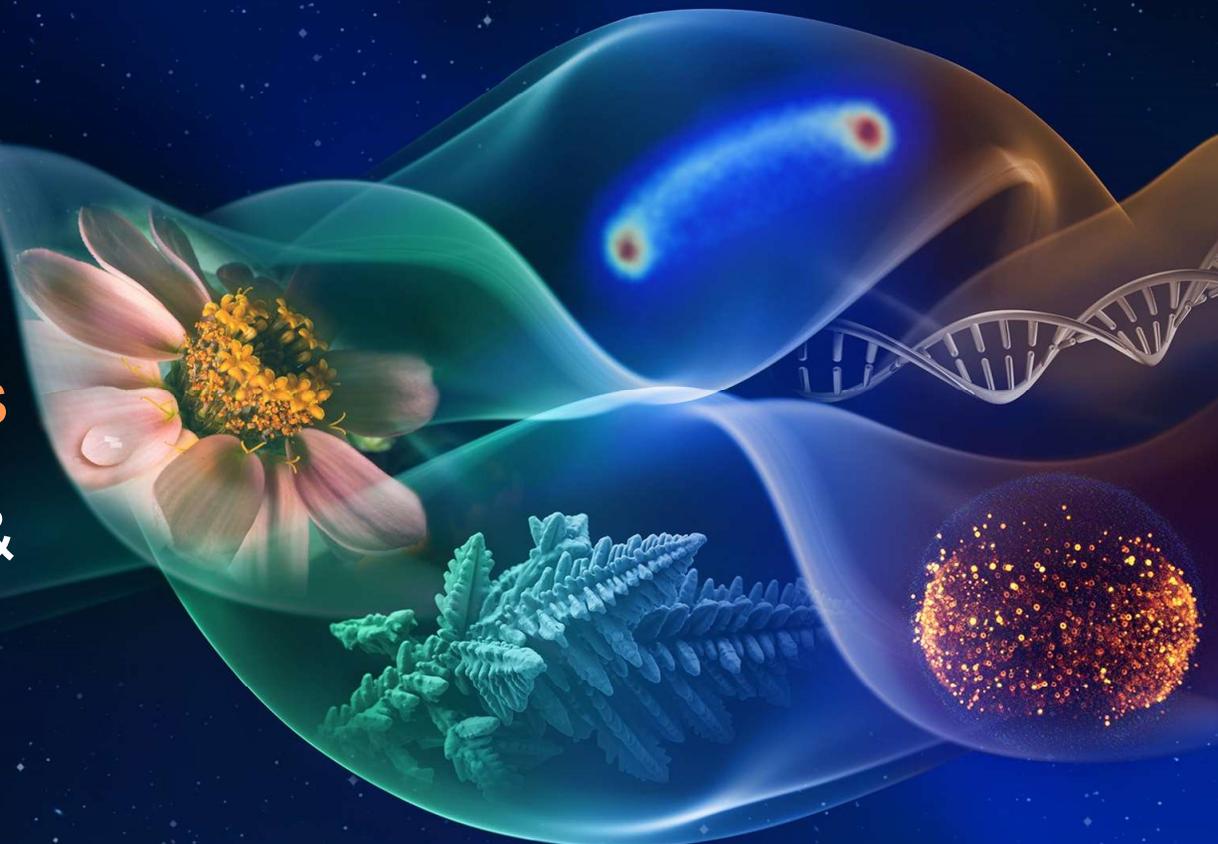
National Aeronautics and
Space Administration



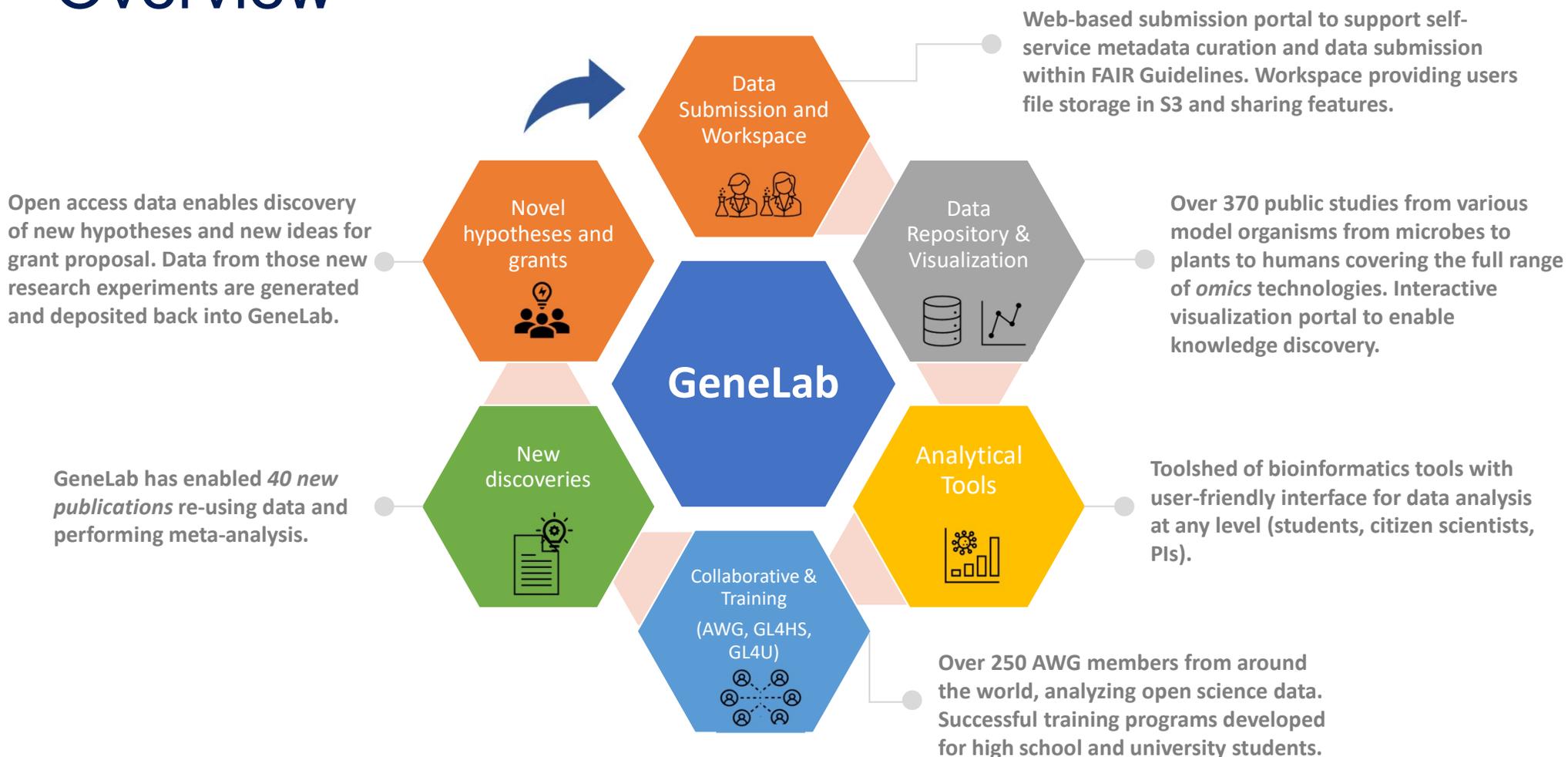
Biological and Physical Sciences

NASEM: GeneLab & Open Science

Sylvain Costes
Open Sciences Data Repository Project
Manager: GeneLab & ALSDA
SCR Branch Chief



Overview



371

Studies

422

Datasets

45

Species

>10

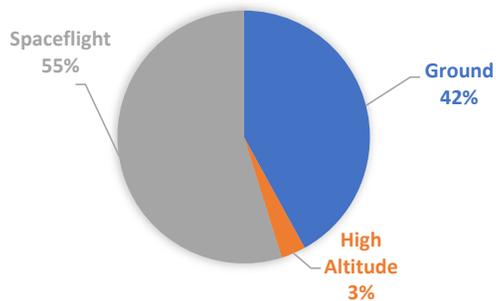
Assays

>135TB

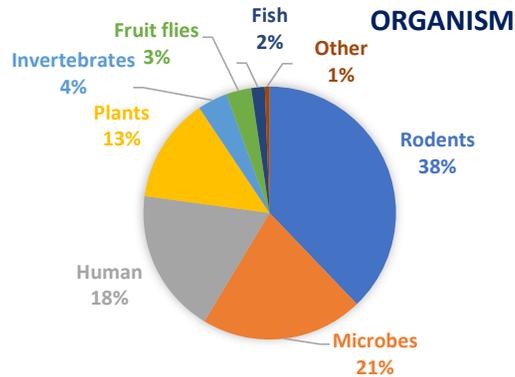
Data

GeneLab Data Repository

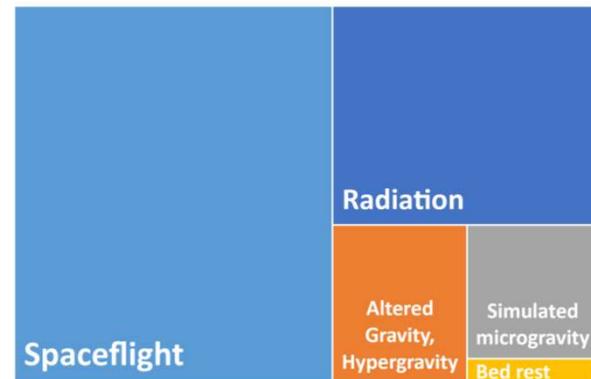
STUDY TYPE



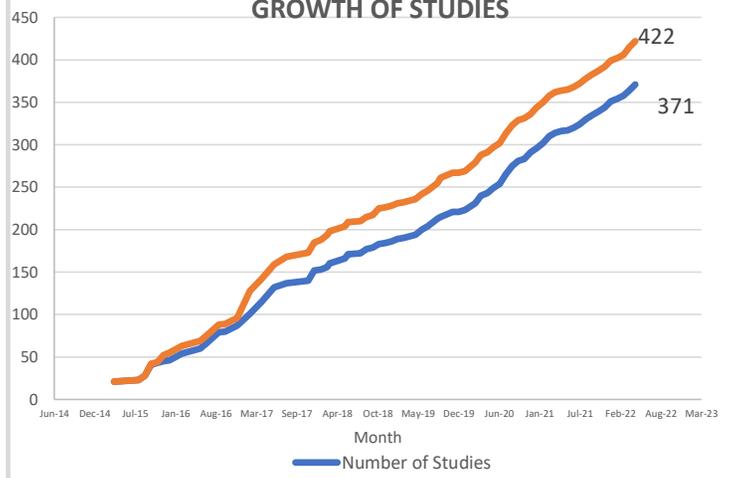
ORGANISM



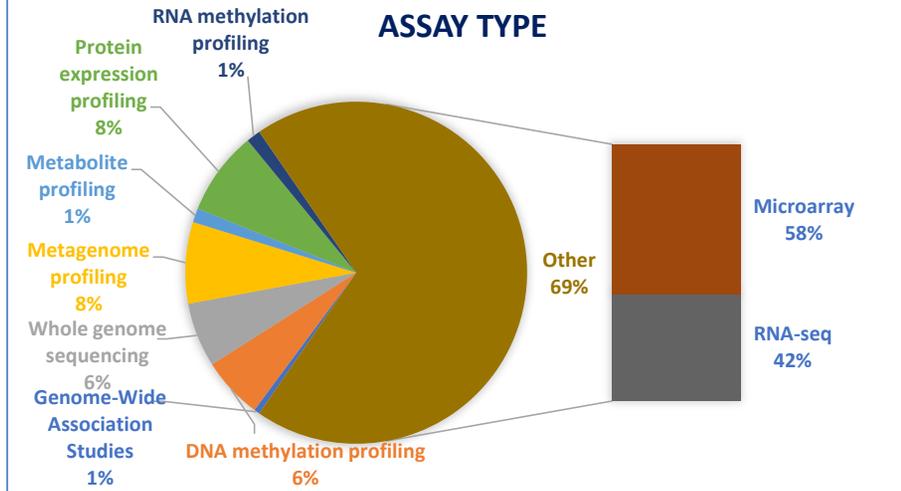
PRIMARY FACTORS



GROWTH OF STUDIES



ASSAY TYPE



Re-use of Data and Enabling New Discoveries

40 enabled publications (10 publications produced by the AWGs) using data available in GeneLab.

Mammalian and Invertebrate Models as Complementary Tools for Gaining Mechanistic Insight on Muscle Responses to Spaceflight

by Thomas Cahill¹, Henry Cope², Joseph J. Bass³, Eliah G. Overbey⁴, Rachel Gilbert^{5,6}, Willian Abraham da Silveira^{1,7}, Amber M. Paul^{5,8,9}, Tejaswini Mishra¹⁰, Raúl Herranz¹¹, Sigrid S. Reinsch⁵, Sylvain V. Costes⁵, Gary Hardiman^{1,12}, Nathaniel J. Szewczyk^{3,13} and Candice G. T. Tahimic^{5,14,*}

An Integrative Network Science and Artificial Intelligence Drug Repurposing Approach for Muscle Atrophy in Spaceflight Microgravity



Vidya Manian¹, Jairo Orozco-Sandoval and Victor Diaz-Martinez

Knowledge Network Embedding of Transcriptomic Data from Spaceflown Mice Uncovers Signs and Symptoms Associated with Terrestrial Diseases

Charlotte A. Nelson¹, Ana Uriarte Acuna^{2,3}, Amber M. Paul^{2,4}, Ryan T. Scott^{2,3}, Atul J. Butte^{5,6}, Egle Cekanaviciute², Sergio E. Baranzini^{1,5,7,*} and Sylvain V. Costes^{2,*}

Cell: The biology of spaceflight package

A coordinated package of 29 scientific papers published in five *Cell Press* journals featuring 9 papers utilizing data or resources in GeneLab. *High impact research highlights:*

- [Comparative Transcriptomics Identifies Neuronal and Metabolic Adaptations to Hyper gravity and Microgravity in *Caenorhabditis elegans*](#), *iScience*
- [Comprehensive Multi-omics Analysis Reveals Mitochondrial Stress as a Central Biological Hub for Spaceflight Impact](#), *Cell*
- [NASA GeneLab RNA-seq consensus pipeline: standardized processing of short-read RNA-seq data](#), *iScience*



Computational and Structural Biotechnology Journal

Volume 19, 2021, Pages 2223-2235



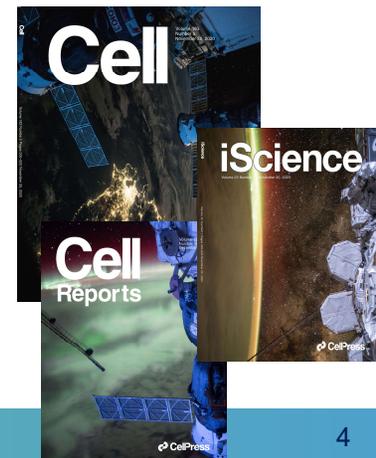
Reanalysis of the Mars500 experiment reveals common gut microbiome alterations in astronauts induced by long-duration confinement

N.J.B. Brereton^{3,8,1}, F.E. Pitre^{3,1}, E. Gonzalez^{5,1}

Rad-Bio-App: a discovery environment for biologists to explore spaceflight-related radiation exposures

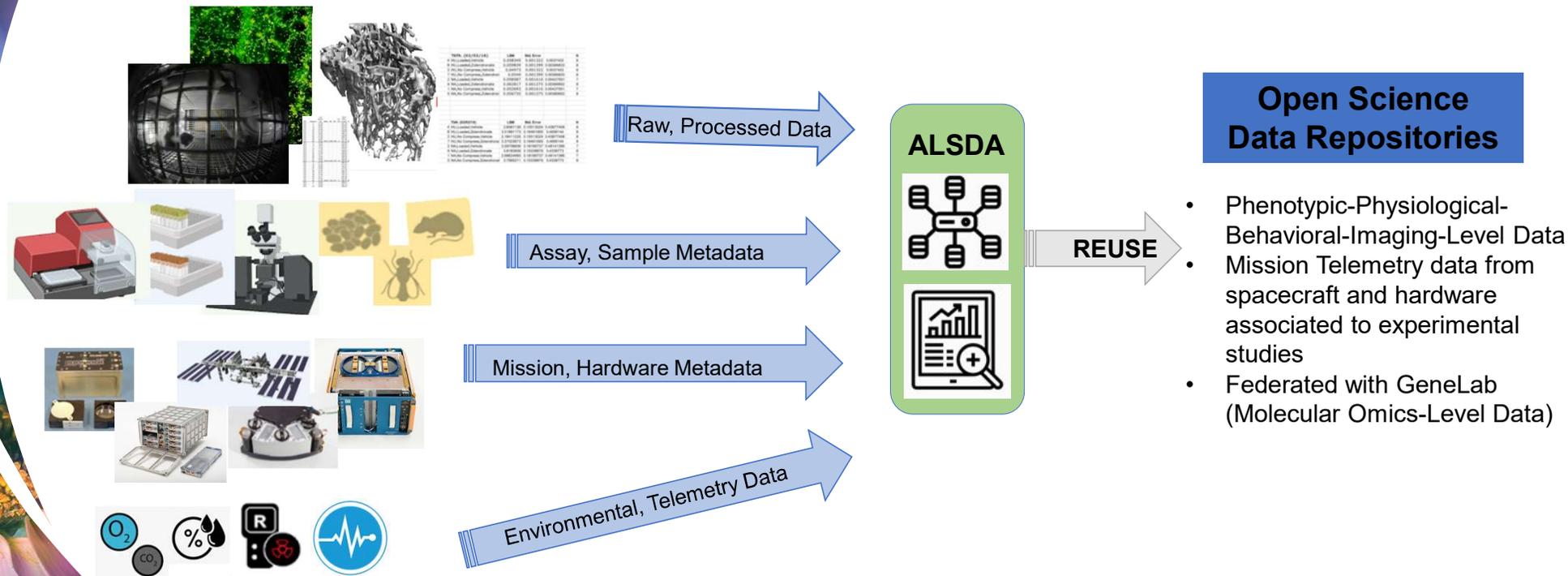
Richard Barker¹, Sylvain V. Costes², Jack Miller³, Samrawit G. Gebre⁴, Jonathan Lombardino⁵ & Simon Gilroy⁶

npj Microgravity 7, Article number: 15 (2021) | [Cite this article](#)



Integrating data beyond *omics*

In 2021, the Ames Life Sciences Data Archive (ALSDA) transitioned to use the GeneLab Data Systems. The goal of this integration was to develop a multi-project submission portal to reduce the burden on PIs submitting data and enabling discovery of both omics and phenotypic data.



(Images from Ronca et al., 2019 <https://doi.org/10.1038/s41598-019-40789-y>; Cells, Bone, RR Habitat NASA-ARC)

Scalability of Data

GeneLab was developed to be scalable – both for data storage and new data types.

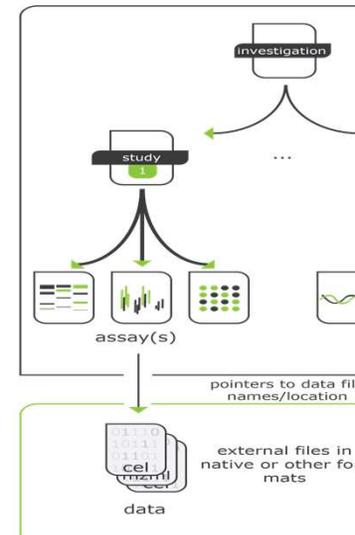
Storage:

- Utilizing cloud for data systems development and data storage
- Cloud provides instant increase in storage
- Allows any data file type



Data:

- Configurable data model to capture any type of experimental data
 - Investigation/Study/Assay (ISA) tab-delimited (TAB) format
- Link multiple assays by sample name
- Not limited to *-omics assays*
- Configure metadata templates to include required fields
 - Qualitative or quantitative measurements (data)
 - Space-relevant fields (for example, radiation dosimetry data per sample)
 - Meets the minimum required information for assays (MIAME, MIxS, MIAPE)
- Machine-readable format – JSON



Omics – new technologies and data types in constant development

In the recent years, new omics technologies have matured such as single cell and spatial omics. Metadata standards were already available and GeneLab was easily able to curate the metadata and organize the files.



GLDS-404

Single cell transcriptional profiling of peripheral blood mononuclear cells (PBMCs) from mice flown on Rodent Research Reference Mission-2 (RRRM-2)

Organisms	Factors	Assay Types	Release Date	Description
Mus musculus	Spaceflight Age	transcription profiling	12-Aug-2021	In the Rodent Research Reference Mission (RRRM-2), forty female C57BL/6NTac mice were flown on the International Space Station. To assess differences in outcomes due to age, twenty 12 week old and two...



GLDS-270

Spatially resolved transcriptional analysis of hearts from mice flown on the RR-3 mission

Organisms	Factors	Assay Types	Release Date	Description
Mus musculus	Spaceflight	transcription profiling	18-Mar-2020	The Rodent Research-3 (RR-3) mission was sponsored by the pharmaceutical company Eli Lilly and Co. and the Center for the Advancement of Science in Space to study the effectiveness of a potential coun...

As new technologies emerge, the GeneLab curation team can create new assay configuration to capture the minimum requirement metadata for each technology. Data standards are provided through standards consortiums and/or scientific community consensus.

Community standards

Standards Consortiums

For omics, many standards for the minimum information about a specific assay experiment exist. For example:

- MIAME - Minimum Information About a Microarray Experiment
- MIAPE – Minimum Information About a Proteomics Experiment
- MlxS – Minimum Information about any (X) Sequence
 - MIGS for genomes
 - MIMS for metagenomes
 - MIMARKS for marker genes
- ENCODE RNAseq experiments

GeneLab uses these standards to ensure data is well curated and complies to the FAIR principles (Findable, Accessible, Interoperable, and Reusable)

Scientific Community Consensus

With the support from our Analysis Working Groups (AWGs), we have developed metadata standards:

- Spaceflight metadata fields by organism type
- Radiation specific metadata fields, including co-developing the Radiation Biology Ontology

ALSDA AWG has contributed their expertise 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 Assay Metadata, Image Configuration for Microscopy	ALSDA Assay Metadata, Image, Outcome Measure Configuration for Same Assay	ALSDA Assay Metadata, Image, Outcome Measure Configuration for Micro-Computed Tomography
File folders will contain: Study Metadata File, Measurement Outcome File, Image of Assay File, Sample Files, Raw Testing Outcome Files	File folders will contain: Study Metadata File, Measurement Outcome File, Image of Assay File, Sample Files, Raw Testing Outcome Files	File folders will contain: Study Metadata File, uCT Assay File, Image of Assay File, Sample Files, Raw Testing Outcome Files
Supporting Literature: [Citation]	Supporting Literature: [Citation]	Supporting Literature: [Citation]
Dimensions (X): [Value]	Dimensions (X): [Value]	Dimensions (X): [Value]
Number of Z sections per image: [Value]	Number of Z sections per image: [Value]	Number of Z sections per image: [Value]
Number of Z segments per image: [Value]	Number of Z segments per image: [Value]	Number of Z segments per image: [Value]
Pixel Type: [Value]	Pixel Type: [Value]	Pixel Type: [Value]
Pixel Size (X): [Value]	Pixel Size (X): [Value]	Pixel Size (X): [Value]
Channel: [Value]	Channel: [Value]	Channel: [Value]
Other behavioral events conducted prior or concurrent after the Assay (see text): [Value]	Other behavioral events conducted prior or concurrent after the Assay (see text): [Value]	Other behavioral events conducted prior or concurrent after the Assay (see text): [Value]
Volume of the region of interest (ROI) in micrometers (µm): [Value]	Volume of the region of interest (ROI) in micrometers (µm): [Value]	Volume of the region of interest (ROI) in micrometers (µm): [Value]
Number of samples per position: [Value]	Number of samples per position: [Value]	Number of samples per position: [Value]
Sample metadata: [Value]	Sample metadata: [Value]	Sample metadata: [Value]
Number of biological markers: [Value]	Number of biological markers: [Value]	Number of biological markers: [Value]
Marker Type: [Value]	Marker Type: [Value]	Marker Type: [Value]
Molecular / Structure label: [Value]	Molecular / Structure label: [Value]	Molecular / Structure label: [Value]
Filteration Algorithm (see text): [Value]	Filteration Algorithm (see text): [Value]	Filteration Algorithm (see text): [Value]
Segmentation Selection Method (see text): [Value]	Segmentation Selection Method (see text): [Value]	Segmentation Selection Method (see text): [Value]
Segmentation Control Method (see text): [Value]	Segmentation Control Method (see text): [Value]	Segmentation Control Method (see text): [Value]

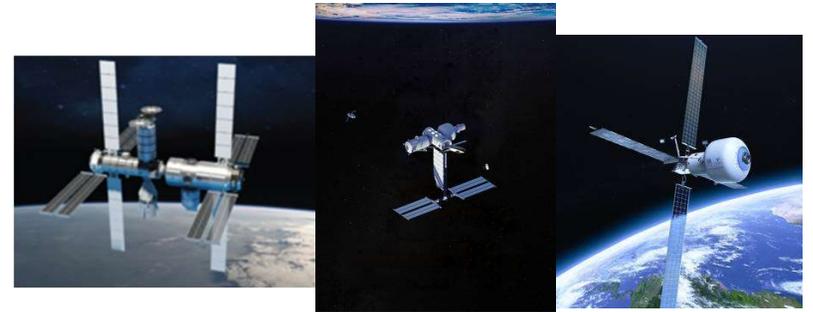
Proprietary data

Currently all the data that is published, is openly accessible worldwide. GeneLab has already in place a mechanism for account-based access, for example our Collaborative workspace. This capability can be expanded in the future for:



Astronaut/Human space-relevant data

- Develop a controlled access data portal
- Create an automatic data access request workflow
- Scientists will be able to translate their findings from model organisms to human outcomes



Commercial Space Industry

- Develop a controlled access data portal and an automatic data access request workflow for “closed” data
- Utilize GeneLab infrastructure to collect and host space-relevant data for biological research

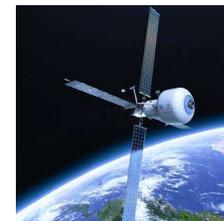
Commercial and International Partners

Open access data in GeneLab, **accelerates the pace of scientific discovery** and thus **enables exploration and development of countermeasures** by using systems biology approaches to identify cross-species relevance, mechanisms, biomarkers and potential health risks for LEO and beyond LEO (Lunar and Mars Exploration).

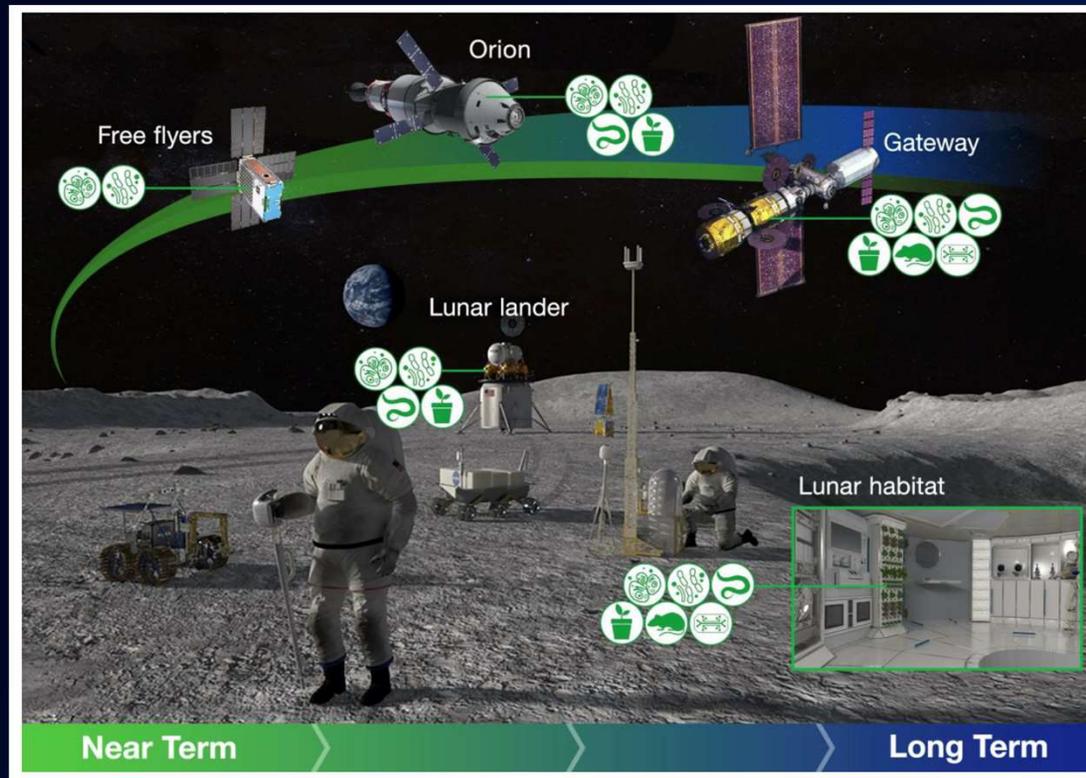
To increase usability and accessibility, space biological research should be available through a common portal. It's important to work closely with our commercial and international partners to ensure data across agencies and platforms are harmonized and comparable. In addition, systems already developed can be utilize not duplicating efforts and costs for data sharing.

For example, GeneLab already hosts datasets from multiple space agencies (ESA, JAXA, CSA) and from CASIS funded experiments.

In order to continue maximizing the return of spaceflight experiments, we encourage commercial and international partners to utilize the GeneLab Data Systems for their research data, both for open and proprietary data.

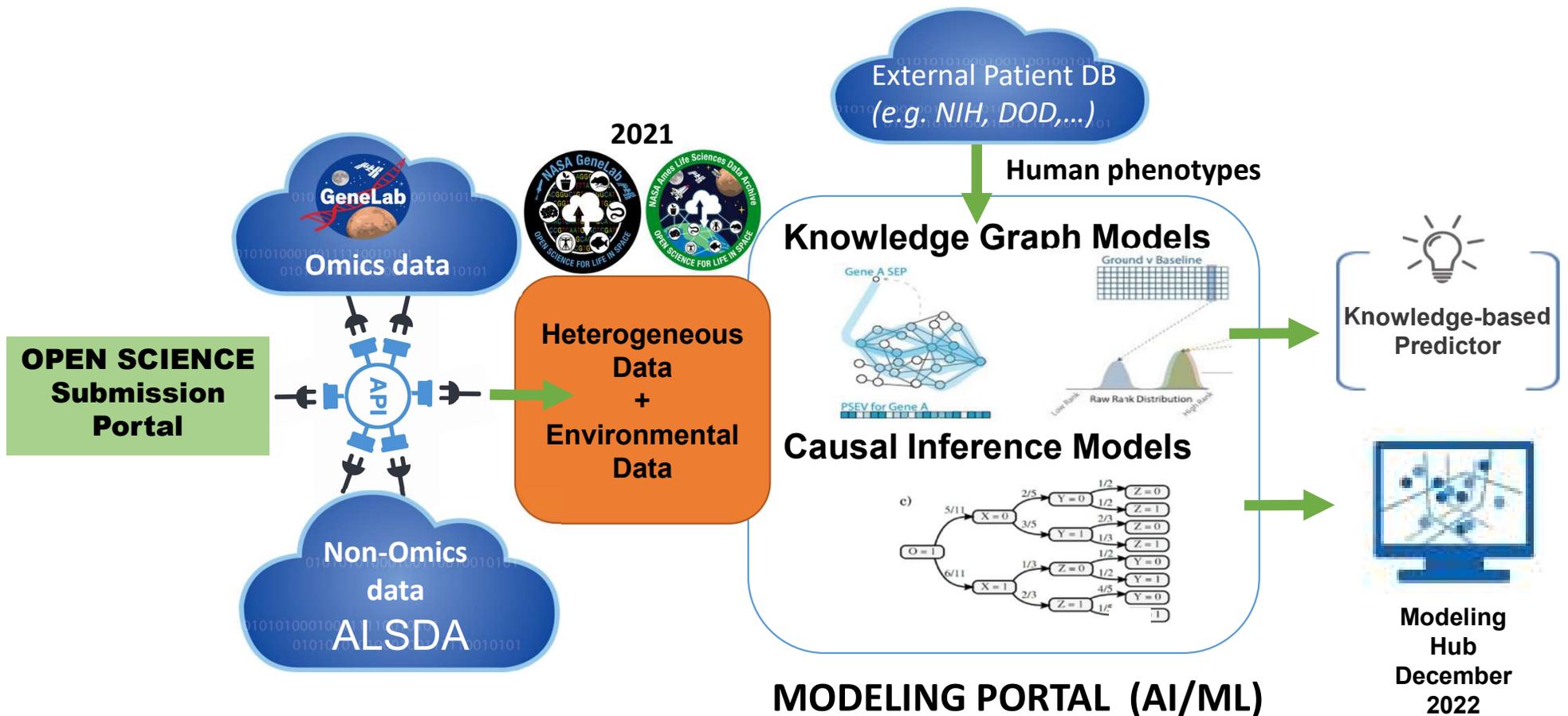


Beyond LEO - Lunar

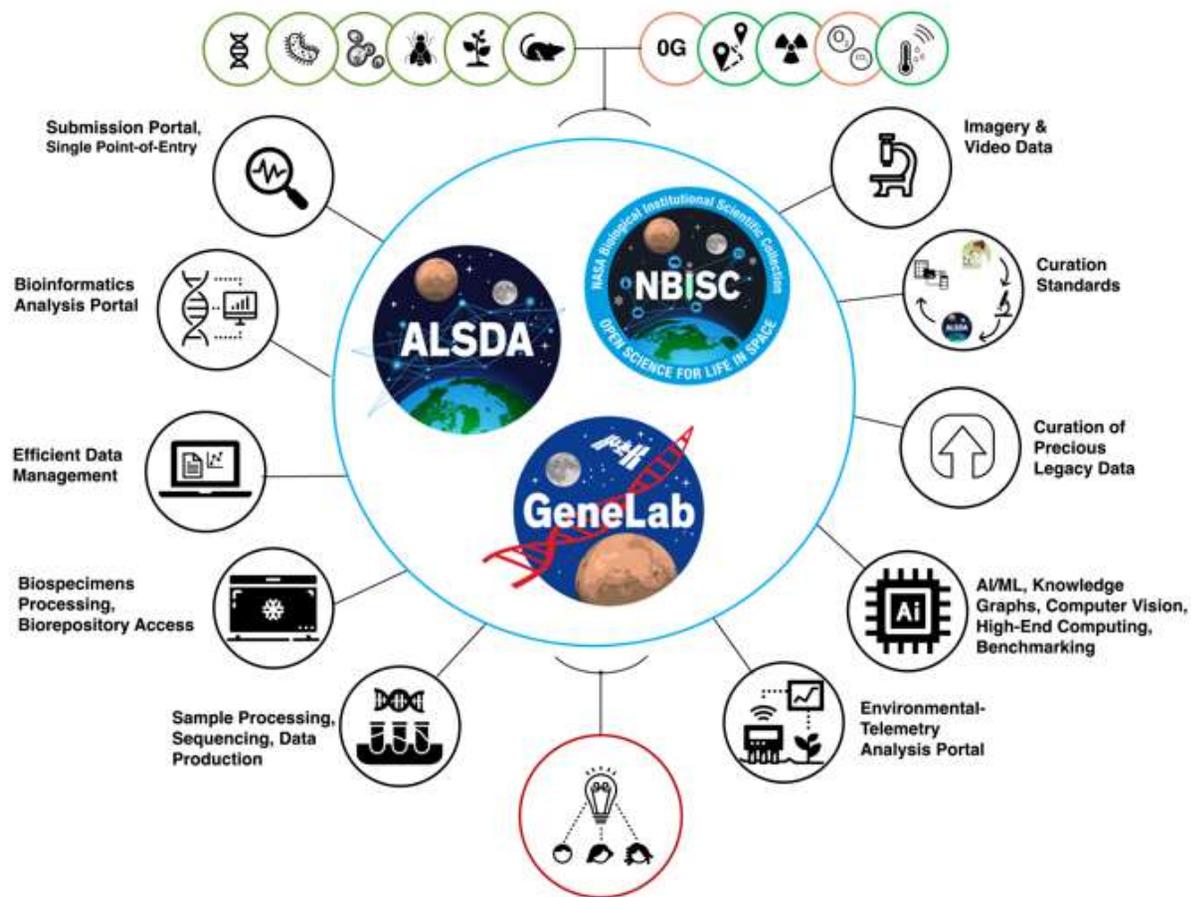


With limited sample return for various platforms, we are preparing new metadata templates and data streams to host and publish data from future lunar experiments and beyond.

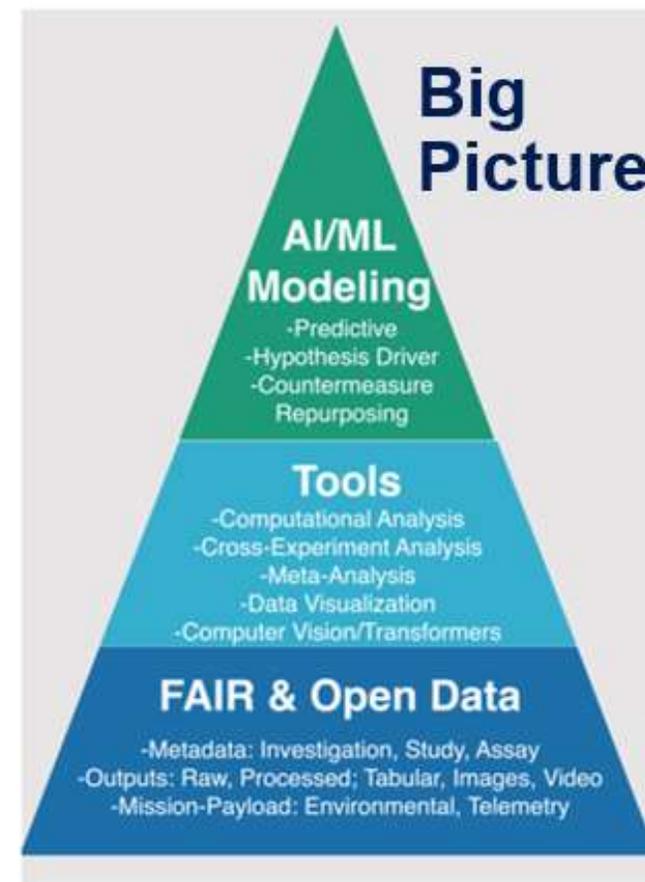
From multiple databases to a knowledge-based system



Open Science Data Repositories



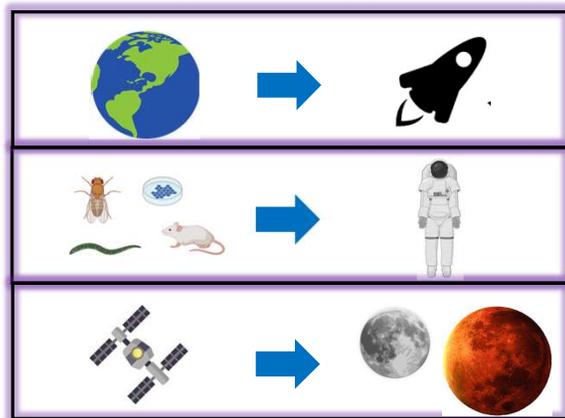
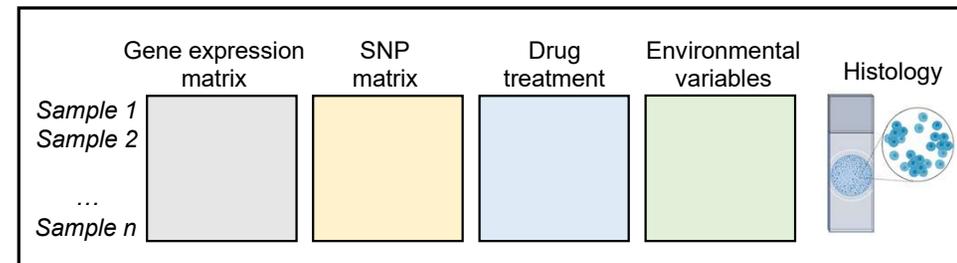
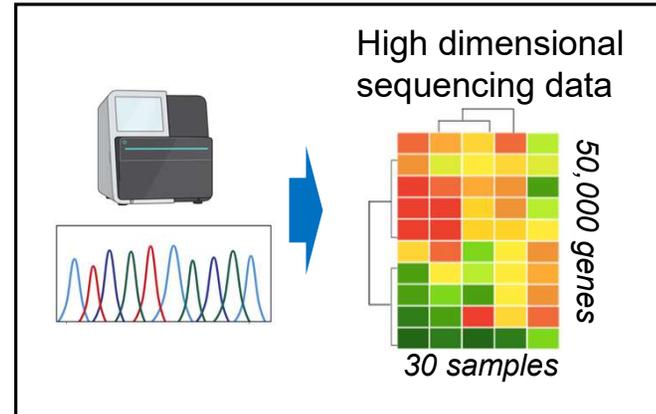
**New Knowledge; Faster Pace;
Inclusive & Diverse Access**



Challenges for Machine Learning in Space Biology

Space Biology Challenges

- Low sample size n
- High dimensionality D
- Heterogeneous data
- Sparse data
- **Out-of-distribution generalization - the testing distribution is unknown and different from the training**
 - (Earth -> space, mouse -> human, short term -> long duration spaceflight)



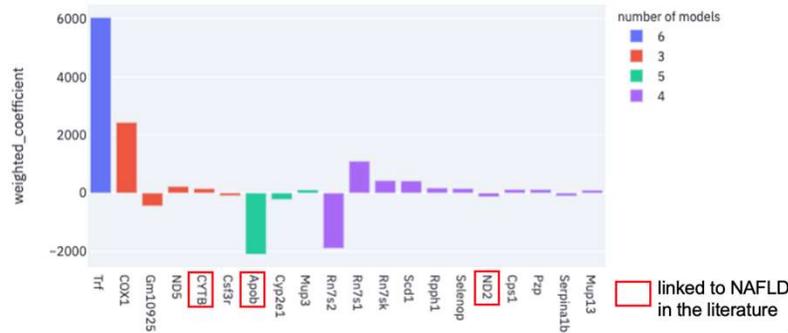
Causal inference in biological systems

Causal Relation Inference and Search Platform (CRISP)

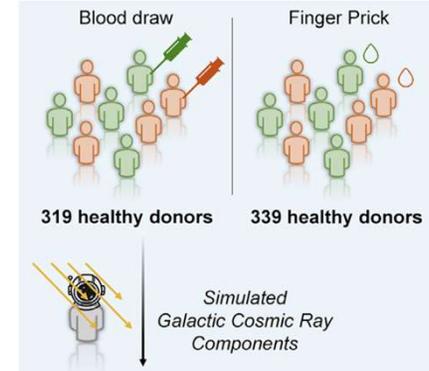


Genes with a causal relationship to liver disease phenotype in spaceflown rodents

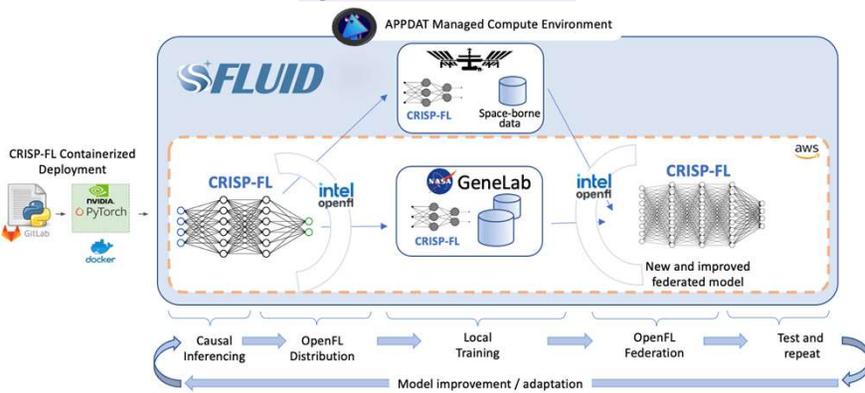
GLDS-47 GLDS-137
GLDS-48 GLDS-168



Genetic variants with a causal relationship to DNA damage and radiation sensitivity in human irradiated immune cells



Federated causal inference between Earth data and spaceborne data



2021 FDL Astronaut Challenge

A novel paradigm for generating human-relevant biomedical insights from observational datasets with limited size by leveraging and augmenting with animal model data

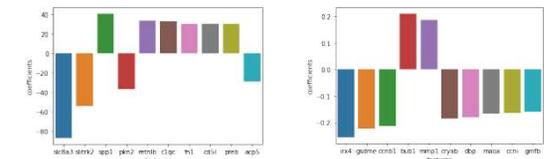
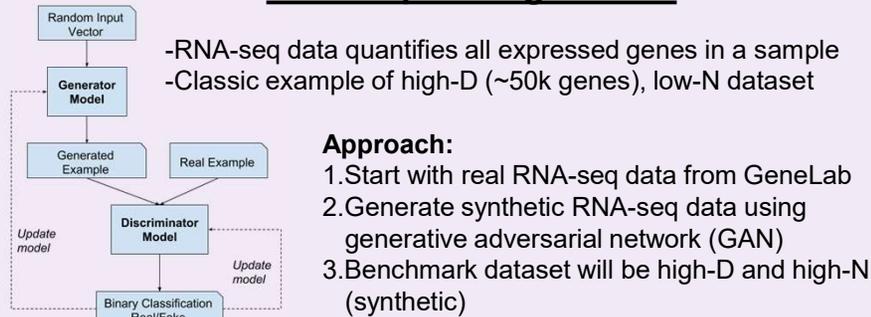


Figure 1: Top-10 IRM features according to feature coefficients with all human and mice gamma-radiation data.

Figure 2: Top-10 IRM features according to feature coefficients with all human and mice heavy-ion radiation data.

AI-readiness and formatting standards: Benchmark datasets for space biology

RNA sequencing dataset



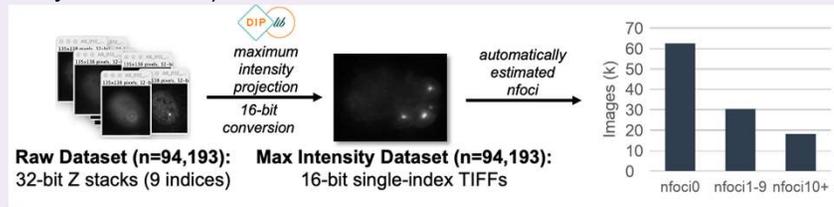
ML applications: e.g. causal inference to identify causal genes with relation to a pathogenic target variable

Microscopy dataset

-Huge amounts of microscopy data in space biology are time consuming to manually annotate through expert review

Approach:

-Mouse radiation exposure microscopy dataset with DNA damage events as distinct foci (PI: Sylvain Costes)



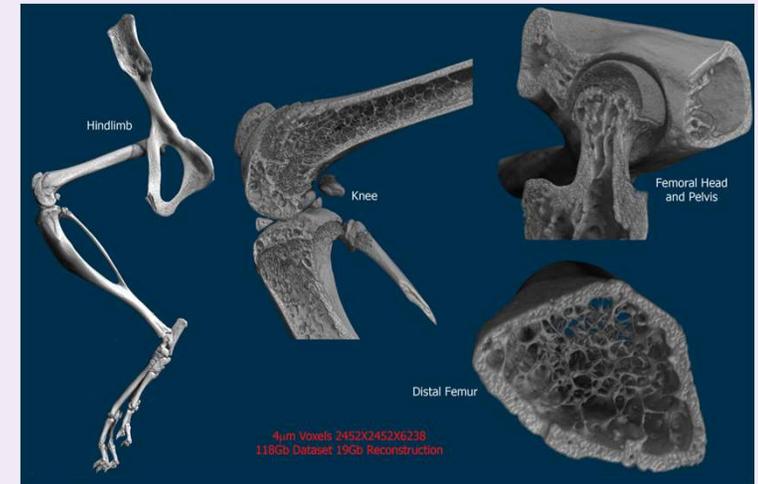
ML applications: e.g. train a self-supervised classifier to detect number of DNA damage foci; or to detect radiation tracks through cells; or to classify images based on the type of radiation.

Bone Computed Tomography (CT) dataset

-CT scan is a key assay in space biology to assess bone changes in spaceflight

Approach:

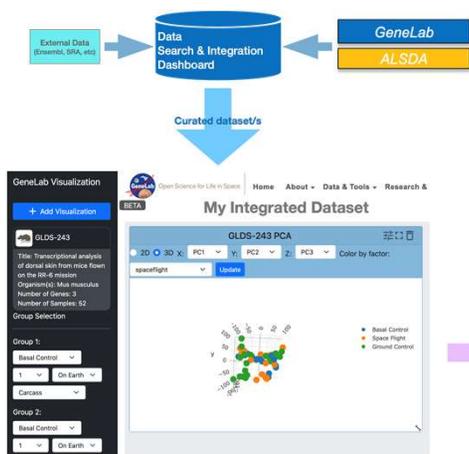
-Mouse hindlimb CT dataset from spaceflight or hindlimb unloading (PI: Eduardo Almeida)



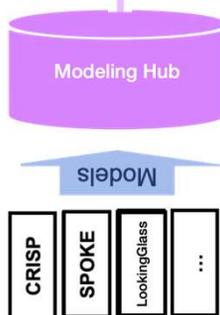
ML applications: e.g. train a classifier to detect the type of stressor based on bone loss, to predict future bone loss when exposed to that stressor.

Space Biology Model Hub and Analysis Portal

Streamlined Data Integration of multiple datasets



Modeling Hub
Automated pipelines for a suite of machine learning and modeling tools



Analysis Portal

Interactive analysis dashboard of modeling and analysis results



Leading the Path to Open-Source Science



Transform to Open Science (TOPS) is a \$40 million* 5-year NASA Science Mission Directorate mission

Objectives:

- ★ Increase understanding & adoption of open science.
- ★ Accelerate major scientific discoveries.
- ★ Broaden participation by historically underrepresented communities.

Goals for 2027:

- ★ 20K earn Open Science Badge
- ★ 5+ major discoveries
- ★ Increase participation of underrepresented groups by 2x



Year of Open Science

*pending appropriations

Training the Next Generation of Scientists



GeneLab for High Schools (GL4HS): A four-week intensive training summer program for rising high school juniors and seniors to learn bioinformatics and computational biology methods and techniques to analyze space omics data.

Learn more and apply at: <https://www.nasa.gov/ames/genelab-for-high-schools>



GeneLab for Colleges/Universities (GL4U): For educators and students to learn how analyze omics data using GeneLab standard pipelines and space-relevant data

Access to course materials: <https://github.com/nasa/GeneLab-Training/tree/main/GL4U>



Space Life Sciences Training Program (SLSTP): Provides undergraduate students entering their junior or senior years, and entering graduate students, with professional experience in space life science disciplines.

Learn more and apply at: <https://www.nasa.gov/ames/research/space-life-sciences-training-program>



Spaceflight Technology, Applications, and Research (STAR): The annual course targets principal investigators (PIs), senior research scientists, and postdoctoral scholars and aims to facilitate their entry to space biology and preparation for conducting spaceflight experiments using NASA and commercial platforms.

Applications open until May 30th: <https://science.nasa.gov/biological-physical/programs/star>



NASA Postdoctoral Program: If you're an early-career or senior scientist, apply to the NASA Postdoctoral Program to help pursue NASA's mission and experience the world's most diverse technology and expertise.

Learn more at: <https://npp.orau.org/>

Near-term Goals

Oct 1st, 2022

Release the Multi-Project
Submission Portal

New Interface and curated
phenotypic data (ALSDA)

FY 2023

Release Modeling Hub and Analysis
Portal

Enable discovery of imaging data
Ease data ingest > 1 TB

FY 2024

Develop smart assist submission
features

Automate data stream from lunar
experiments

Participate in SMD-Source Science Initiatives – Present opportunities to the BPS scientific community

Continue to ingest and host experimental and mission telemetry from space-relevant experiments

Pending yearly budget

Thank you!



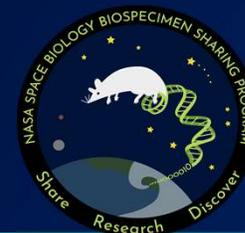
Twitter: @NASAGeneLab
<https://genelab.nasa.gov>



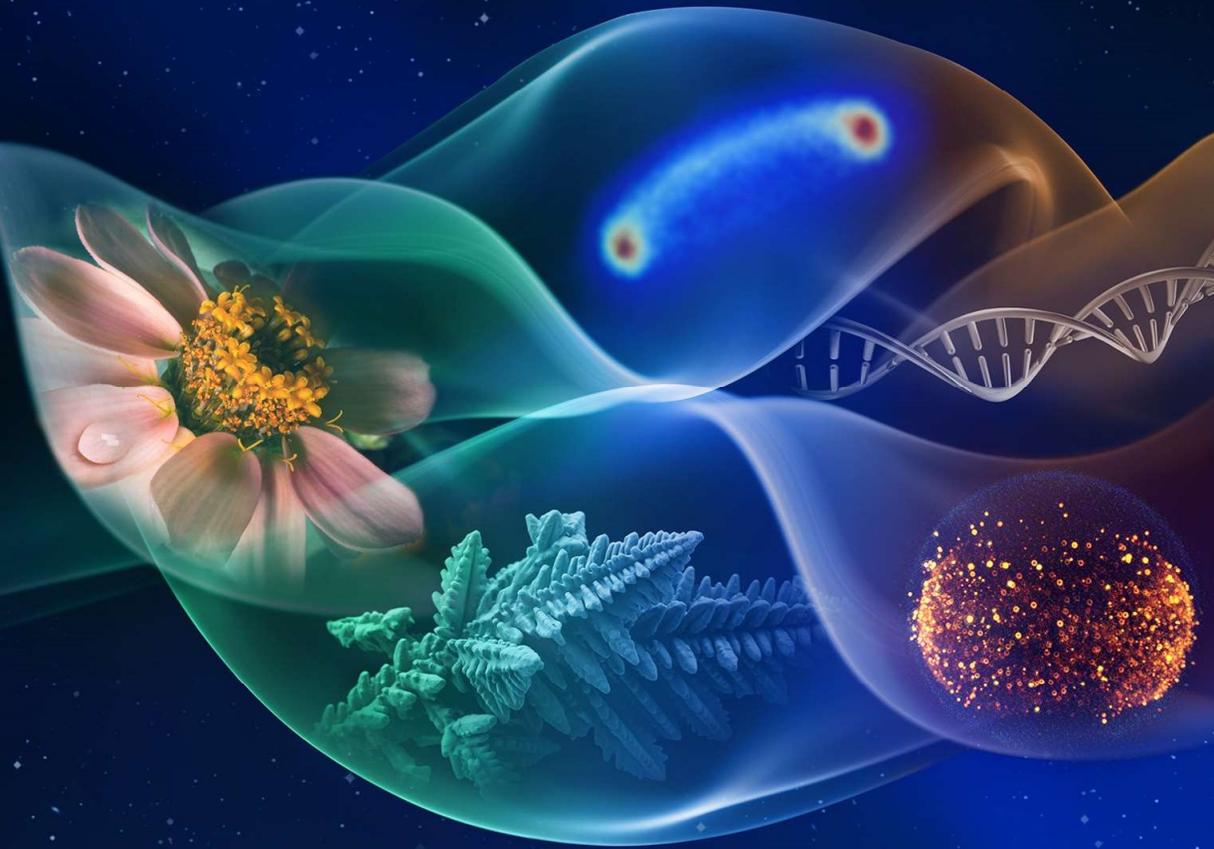
GeneLab is funded by the NASA Space Biology program within the NASA Science Mission Directorate's (SMD) Biological and Physical Sciences (BPS) Division



Open Science for Life in Space is funded by the NASA Space Biology program and NASA Human Research Program



BACK UP



NASA Biological Open Science Resources

Biospecimen Sharing Program (BSP)



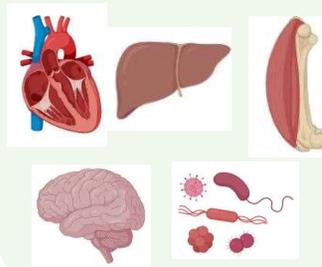
- *Dissects and preserves* rodent tissues from Flight and Ground investigations
- *Coordinates* internal tissue sharing



NASA Biological Institutional Scientific Collection (NBISC)



- *Collection* of non-human specimens and space microbial culture



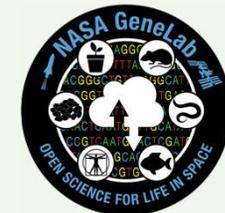
Ames Life Sciences Data Archive (ALSDA)



- *Collects and curates physiological, mission, project and imaging data*



GeneLab (GL)



- *Collects and curates omics data*

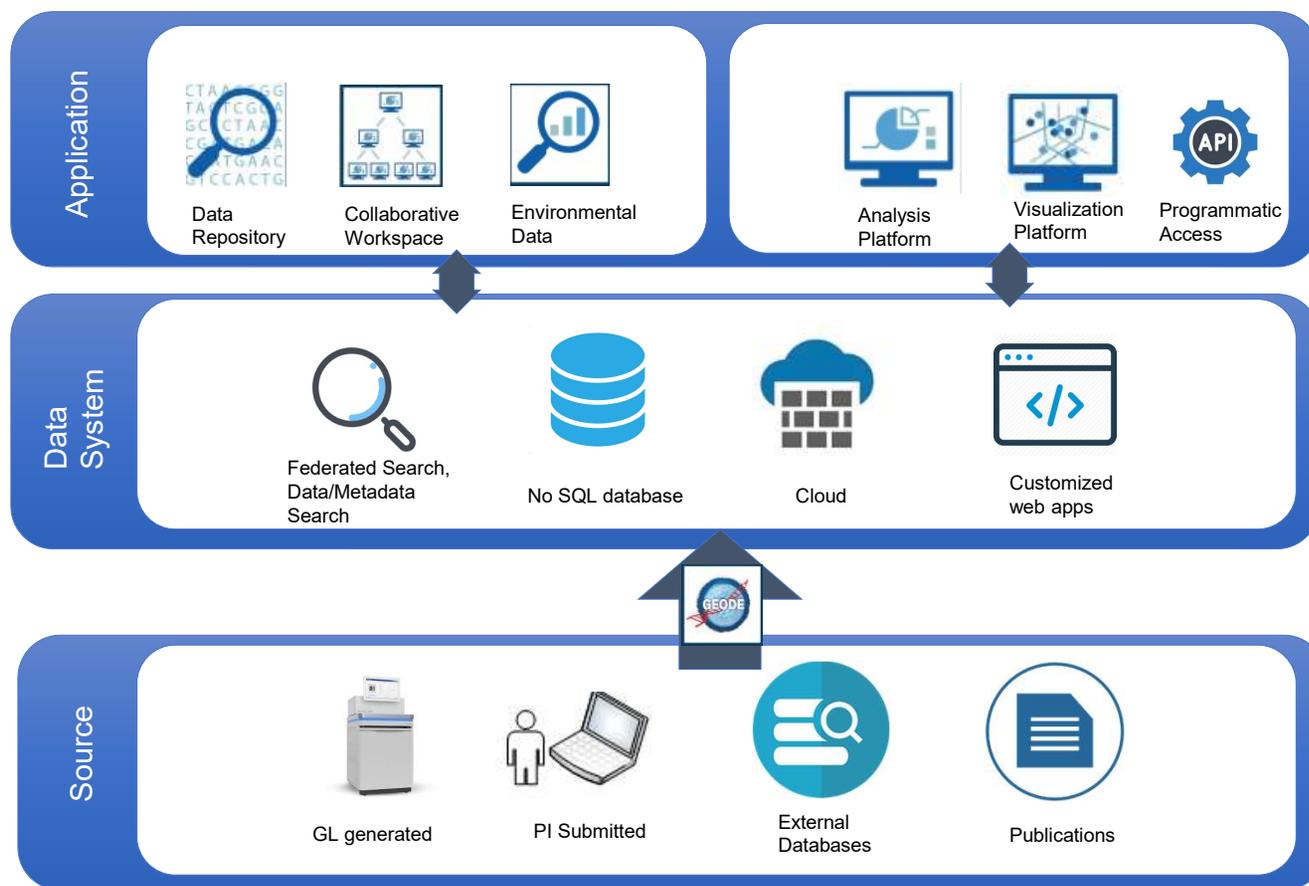


NASA BPS Open Science Enterprise Solution: genelab.nasa.gov

- Open access data
- **FAIR (Findable, Accessible, Interoperable, Reusable)**
- Controlled access tools
- API - internal and external
- User Friendly Interface
- Tutorials
- Self-service Submission Portal

- Federated search – GEO, PRIDE, MG-RAST, **ALSDA**
- Database & Cloud – Scalable, easy access, fast
- Web apps
 - Data Access & Management
 - Security
 - Operation
 - Governance and Integration
- Open Source software – *no maintenance cost for software*

- Multiple data sources
 - Standard metadata organization
 - Open file formats



Omics Data Generation

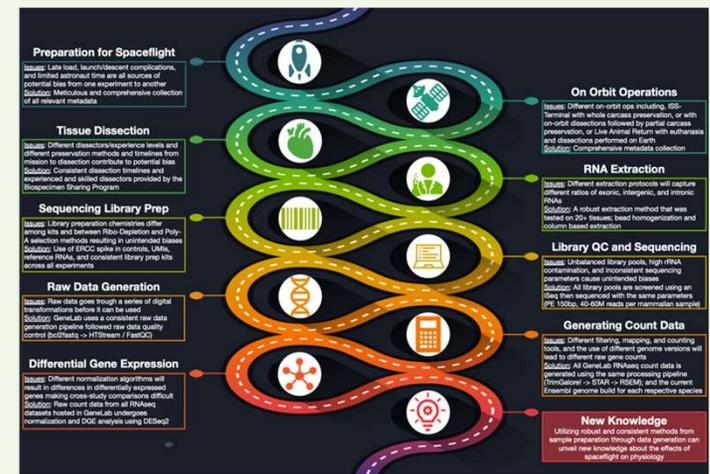
Maximizing spaceflight experiments by generating omics data without any proprietary time for open science.

- Maximizing access to spaceflight data through tissue sharing efforts.
- Upon curation, these data are released *immediately* to the public through the GeneLab data system.
- To date, **59 studies** are available including data from: Rodent Research missions and Plant experiments. Assays include:
 - Transcriptomics
 - Bulk RNASeq
 - Single Cell RNASeq
 - Spatial Transcriptomics
 - Epigenomics
 - Proteomics
- Enabling meta-analysis publications



Standardizing Sample Processing and Sequencing for Spaceflight Samples

- GeneLab also works with NASA-funded principal investigators (PI's) to generate proprietary data using GeneLab standard assays.
- Optimized processing protocols and high-quality data outputs
- Data Processing services
- Request quote: <https://genelab.nasa.gov/form/genelab-sequencing-services>



GeneLab Power Users

GeneLab Analysis Working Groups (AWGs) consist of 250+ scientists from multiple space agencies, international institutions, and industry. Scientists meet monthly with each group to analyze data in the GeneLab repository. We invite you to join - <https://genelab.nasa.gov/awg/join!>

ANIMAL

68 members

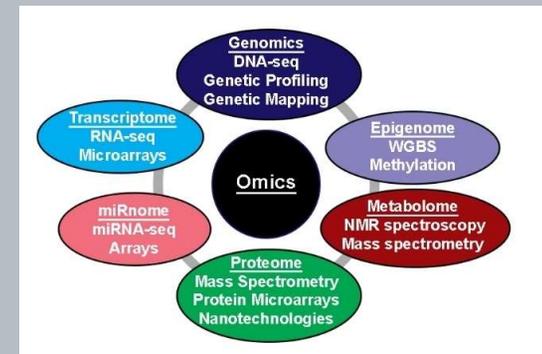
Facilitates the use of omics in understanding basic mechanisms by which animals and constituent tissues and cells adapt to the spaceflight environment.



MULTI-OMICS

128 members

Interactions between the different omics to provide complete understanding of the entire system being studied.



PLANTS

58 members

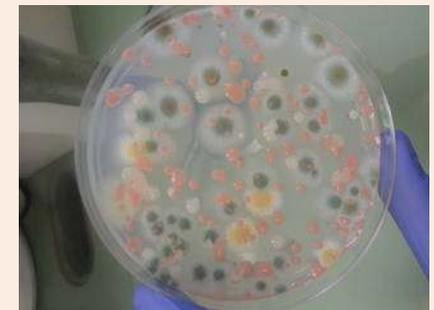
Share and discuss the latest developments in **Astrobotany** – the discipline of botany concerned with interactions between plant biology and space environment.

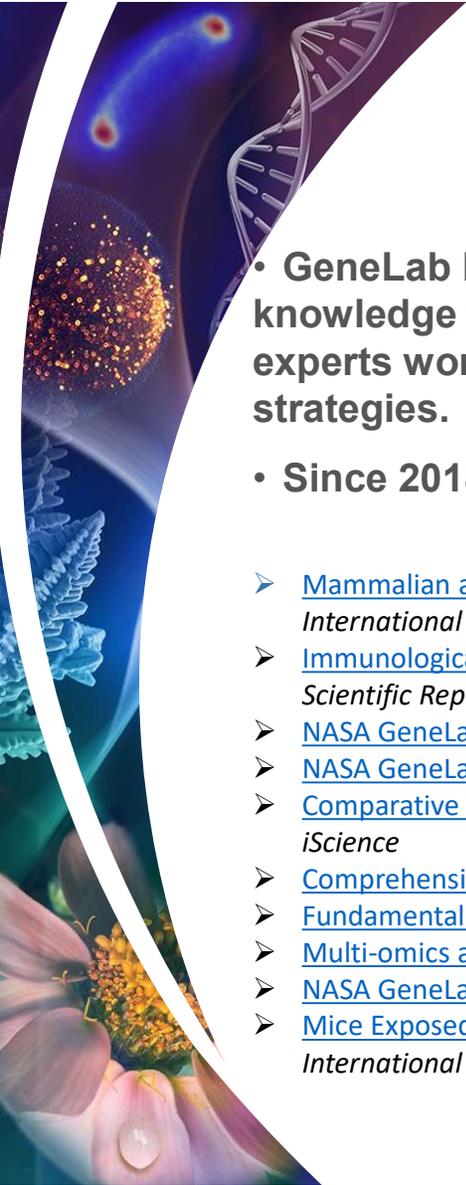


MICROBES

50 members

Focuses on analyzing microbial datasets within GeneLab that includes gene-expression, proteomic, metabolomic and environmental metagenomic datasets.





Accelerating the pace of scientific discovery through open data

- GeneLab has established the AWGs with the goal of maximizing the generation of new knowledge from these rare and complex spaceflight datasets. With over 250+ scientists and experts worldwide the four working groups focus Animal, Plants, Microbe and Multi-omics strategies.
- Since 2018, the AWGs have published 10 peer-reviewed publication re-using data in GeneLab:
 - [Mammalian and Invertebrate Models as Complementary Tools for Gaining Mechanistic Insight on Muscle Responses to Spaceflight](#), *International Journal of Molecular Sciences*
 - [Immunological and hematological outcomes following protracted low dose/low dose rate ionizing radiation and simulated microgravity](#), *Scientific Reports*
 - [NASA GeneLab RNA-seq consensus pipeline: standardized processing of short-read RNA-seq data](#), *iScience*
 - [NASA GeneLab RNA-seq consensus pipeline: standardized processing of short-read RNA-seq data](#), *Cell System*
 - [Comparative Transcriptomics Identifies Neuronal and Metabolic Adaptations to Hypergravity and Microgravity in *Caenorhabditis elegans*](#), *iScience*
 - [Comprehensive Multi-omics Analysis Reveals Mitochondrial Stress as a Central Biological Hub for Spaceflight Impact](#), *Cell*
 - [Fundamental Biological Features of Spaceflight: Advancing the Field to Enable Deep-Space Exploration](#), *Cell*
 - [Multi-omics analysis of multiple missions to space reveal a theme of lipid dysregulation in mouse liver](#), *Science Reports*
 - [NASA GeneLab Platform Utilized for Biological Response to Space Radiation in Animal Models](#), *Cancers*
 - [Mice Exposed to Combined Chronic Low-Dose Irradiation and Modeled Microgravity Develop Long-Term Neurological Sequelae](#), *International Journal of Molecular Sciences*