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Fiscal Year:	FY 2021	Task Last Updated:	FY 12/05/2020
PI Name:	Carr, Christopher Sc.D.		
Project Title:	Enterococci Evolution in Space: Environmental Adaptations, Antibiotic Resistance, and Clinical Implications		
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Division Name:	Space Biology		
Program/Discipline:			
Program/Discipline Element/Subdiscipline:			
Joint Agency Name:		TechPort:	No
Human Research Program Elements:	None		
Human Research Program Risks:	None		
Space Biology Element:	(1) Cell & Molecular Biology(2) Microbiology		
Space Biology Cross-Element Discipline:	None		
Space Biology Special Category:	(1) Translational (Countermeasure) Po	tential	
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Comments:			
Project Type:	FLIGHT	Solicitation / Funding Source:	2016-17 Space Biology (ROSBio) NNH16ZTT001N-FG. App G: Flight and Ground Space Biology Research
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No. of PhD Candidates:		No. of Master' Degrees:	
No. of Master's Candidates:		No. of Bachelor's Degrees:	
No. of Bachelor's Candidates:		Monitoring Center:	NASA ARC
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Flight Program:			
Flight Assignment:			
Key Personnel Changes/Previous PI:			
COI Name (Institution):	Burton, Aaron Ph.D. (NASA Johnson Space Center) Gilmore, Michael Ph.D. (Massachusetts Eye And Ear Infirmary & Physician Staff, Inc.) Wallace, Sarah Ph.D. (NASA Johnson Space Center)		
Grant/Contract No.:	80NSSC21K0234		
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Enterococci are gram-positive bacteria that originated when our ancient animal ancestors emerged from the oceans to live on land, and brought their gut flora with them. Enterococcus faecalis (EF) and Enterococcus faecium, are common human commensals and harbor multidrug resistance. Both have been previously isolated on the International Space Station (ISS). Likely as a consequence of their evolutionary origins, enterococci show remarkable stress resistance within, but also outside, their human hosts. Their antibiotic resistance, coupled with tolerance to desiccation, starvation, and disinfection, make them potent pathogens in the built environment, and a risk to crew health during space missions. Specific Aims: Here we propose a three-year Principal Investigator (PI) team research program, including flight components, to: 1) Characterize the frequency and genomic identity of antibiotic resistant organisms, including enterococci, on the ISS; 2) Assess the evolutionary selective pressure of the space environment (microgravity, space radiation) using EF as a model system; and 3) Characterize the natural evolutionary history of EF on Earth and in space to reveal mechanisms of microbial adaptation including natural selection, and refine crew health risk assessment and countermeasures for future space missions. Each aim is addressed by a PI-led investigation.

Genomic Enumeration of Antibiotic Resistance in Space (GEARS): Led by Co-PIs Burton and Wallace (NASA), this flight experiment will leverage existing ISS facilities to enable detection, species-level identification (via nanopore sequencing), and isolate selection (for Earth return and characterization), complementing ongoing routine microbial monitoring by focusing on the clinically-relevant subset of antibiotic resistant microbes.

Enterococcus Growth Advantage on ISS via Tn-seq (EnteroGAIT): Led by Co-PI Gilmore (Massachusetts Eye And Ear Infirmary, MEEI), this flight experiment will utilize the Bioculture System, currently undergoing validation on the ISS, to measure the selective pressure of the space environment on a defined microbial population: a library of EF mutants, created by transposon insertional mutagenesis, will undergo several rounds of growth and starvation on ISS with a synchronous ground control; selection is measured for each gene knockout by sequencing (Tn-Seq) and occurs on timescales far shorter than natural or experimental evolution.

Adaptation and Evolution of Resilient Enterococcus in Space (AERES): Led by PI Carr (Georgia Tech), this ground-based data and experiment-driven study will use an extensive collection of existing (>500) and newly generated (~100) whole genomes of Earth and space-derived enterococci isolates to characterize horizontal gene transfer, mutation, and implications for community-acquired infection in space and on Earth. Genetic techniques will be used to establish causality.

Significance and Relevance: The proposed work will improve our fundamental understanding of microbial adaptation to the space environment and crew health risks. Our work directly addresses the solicitation research foci A.3.1.a "a. Conduct long-term, multigenerational studies of microbes using the ISS or other appropriate flight platforms as a Microbial Observatory to study and understand the population and community dynamics of the microbes that inhabit that unique environment" (primary) and A.3.1.b "b. Determine the influence of spaceflight on defined microbial populations and communities. Space Biology studies will determine the effects of spaceflight on dynamics of microbes in mono or mixed cultures with respect to cell processes (including virulence and antibiotic resistance, evolution, biofilm formation, and community development)" (secondary) and the 2011 Decadal Survey recommendation to use the ISS as a Microbial Observatory (P1). Furthermore, our work addresses multiple Human Research Roadmap (HRR) gaps (MICRO-1 to MICRO-5 and Med01).

Rationale for HRP Directed Research:

Research Impact/Earth Benefits:

Task Description:

Task Progress: New project for FY2021.

Bibliography Type: Description: (Last Updated: 09/15/2023)