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Boston Children's Hospital. 10) Added Gervasio Batista, PhD to role of Post-Doc in Dr. Hensch's laboratory at Harvard Boston Children's Hospital.

Basner, Mathias M.D., Ph.D. (University of Pennsylvania) Bilker, Warren Ph.D. (University of Pennsylvania) Chouker, Alexander M.D. (University of Munich) Elliott, Mark Ph.D. (University of Pennsylvania) Feiveson, Alan Ph.D. (NASA Johnson Space Center) Flanagan, Shawn Ph.D. (University of Pittsburgh) Gehrman, Philp Ph.D. (University of Pennsylvania)

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COI Name (Institution): Kuehn, Simone Ph.D. (University Clinic Hamburg-Eppendorf)
Landon, Lauren Ph.D. (KBR/NASA Johnson Space Center)

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Bell, Suzanne T Ph.D. (Behavioral Health & Performance Lab at KBR/NASA JSC )
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Grant/Contract No.:

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**Performance Goal No.:** 

**Performance Goal Text:** 

Task Description:

NASA's vision for successful long-duration exploration missions (LDEM) depends on optimizing human performance, adaptability, and resiliency to reduce individual and crew behavioral risks. To date, the major emphasis in optimizing astronauts for their tolerance to prolonged spaceflight has involved human health and performance countermeasures as well as technologies and tools to ensure safety during exploration. However, considerable evidence suggests that there are individual differences among astronauts in their vulnerabilities to the various stressors of spaceflight. The goal of the proposed NSCOR (NASA Specialized Center of Research) is to obtain novel information that will help identify individuals who are resilient to the stressors of prolonged human spaceflight, thereby ensuring successful completion of exploration missions and the preservation of astronaut health over the life of the astronaut. This NSCOR project leverages the NIMH (National Instsitute of Mental Health) Research Domain Criteria (RDoC) heuristic framework to conduct experimental studies to identify biological domains (molecular, circuitry, physiology) and behavioral domains that relate to individual adaptation and resiliency (as well as behavioral vulnerability) in spaceflight-relevant isolated confined and extreme environments (ICC and ICE). The NSCOR focuses specifically on differences among astronauts in their tolerance of and adaptability to simulated conditions of prolonged spaceflight that impact behavioral health and performance. The NSCOR will provide novel information on the extent to which behavioral and biological factors can be identified that predict astronauts who can maintain positive mood, proactive social processes, a high level of performance and personal well-being, while coping with confinement, meaningless work, limited social support, and living in the extreme environmental conditions of space. By utilizing the RDoC framework, three different human confinement analogs and an animal model, the NSCOR will generate data converging on biomarkers of neurobehavioral and neurobiological resilience to the spaceflight conditions. Such a discovery will help in selecting astronauts most likely to maintain human health and performance during long-duration exploration missions.

## Rationale for HRP Directed Research:

**Research Impact/Earth Benefits:** 

This project will benefit the spaceflight community, specifically humans involved in spaceflight, by providing information that will help characterize the three less well-understood NIMH RDoC domains related to positive valence, negative valence, and social processes as they relate to performance, adaptation, and resilience of individuals living and working in ICC/ICE environments. We will identify predictive indicators and biomarkers for resilience and adaptation in individuals to aid in selection and individualized countermeasure development with the goal to maintain and optimize performance capability and behavioral health during Long Duration Exploration Missions.

Regarding the human analogs, the investigator team has continued to hold regular electronic virtual meetings (N=18 over this reporting period) to discuss and resolve important issues related to data acquisition, updated literature on resilience and potential biomarkers, data storage, quality control and analyses, and the resumption of analog research. This enabled the team to update its scheduled study runs and refine its list of biological biomarkers and questionnaires. These meetings facilitated defining resilience with the paired comparison methodology, as well as the development of a pipeline for data management, data harmonization, and data analysis to ensure consistency across NSCOR analog sites for heart rate variability analyses.

The COVID-19 pandemic resulted in a mandated cessation of scheduled NSCOR data acquisition at the Isolation and Confinement Analog Research Unit for Spaceflight (ICARUS) facility at the University of Pennsylvania. Data acquisition is scheduled to recommence this year, if the University of Pennsylvania Perelman School of Medicine approves the updated research resumption plan. The team at Penn has continued to work with the statistician, Dr. Warren Bilker, to create a resilience score for the NSCOR across analog sites. Data acquisition for Campaign 6 in Human Exploration Research Analog (HERA) is postponed due to the COVID-19 pandemic. Campaign 6 is currently scheduled to begin this year. Data acquisition for the second 14 month mission in Neumayer is complete.

During the past year in the mouse model, Dr. Hensch and colleagues identified sex-specific consequences of early life adversity in an adult mouse model. These results may impact the selection of candidates or optimal environments for long-term spaceflight confinement stress in humans based on their adverse childhood experience (ACE) scores. To capture the true complexity of social behaviors that occur in groups, they established a Social Network Analysis (SNA) approach to define objective parameters associated with sociability and its plasticity by sex. This analytical approach can identify, for example, network participants who are more effective teammates or more likely to generate new social

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	relationships. Anticipated baseline behavior was further compared to novel and established molecular biomarkers to be assessed in peripheral samples (e.g., urine, blood), including autoantibodies and oxidative stress markers. Importantly, the ability to overcome impairments of sociability with acute antioxidant (or other) treatments can now be tested in animal models.
Bibliography Type:	Description: (Last Updated: 03/24/2024)