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Fiscal Year:	FY 2023	Task Last Updated:	FY 12/08/2022
PI Name:	Lee, Stuart M.C. Ph.D.		
Project Title:	Determining the Dose Response Profile of the Headward Fluid Shift during Varying Gravity Levels		
Division Name:	Human Research		
Program/Discipline:			
Program/Discipline Element/Subdiscipline:			
Joint Agency Name:		TechPort:	No
<b>Human Research Program Elements:</b>	(1) <b>HHC</b> :Human Health Countermeasures		
Human Research Program Risks:	(1) Cardiovascular: Risk of Cardiovascular Adaptations Contributing to Adverse Mission Performance and Health Outcomes		
Space Biology Element:	None		
Space Biology Cross-Element Discipline:	None		
Space Biology Special Category:	None		
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Zip Code:	77058-2749	<b>Congressional District:</b>	36
Comments:			
Project Type:	GROUND	<b>Solicitation / Funding Source:</b>	Directed Research
Start Date:	03/01/2021	End Date:	06/30/2024
No. of Post Docs:	0	No. of PhD Degrees:	0
No. of PhD Candidates:	0	No. of Master' Degrees:	0
No. of Master's Candidates:	0	No. of Bachelor's Degrees:	0
No. of Bachelor's Candidates:	0	<b>Monitoring Center:</b>	NASA JSC
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Flight Program:			
Flight Assignment:	NOTE: End date changed to 06/30/2024 per C. I	Ribeiro/HHC (Ed., 10/5/23).	
Key Personnel Changes/Previous PI:			
COI Name (Institution):	Martin, David M.S. (KBR/NASA Johnson Space Center ) Laurie, Steven Ph.D. (KBR/NASA Johnson Space Center ) Macias, Brandon Ph.D. (NASA Johnson Space Center ) Marshall-Goebel, Karina Ph.D. (KBR/NASA Johnson Space Center ) Young, Millennia Ph.D. (NASA Johnson Space Center ) Besnard, Stephane M.D., Ph.D. (University Hospital of Caen, France )		
Grant/Contract No.:	Directed Research		
Performance Goal No.:			
Performance Goal Text:			

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Long-duration stays in weightlessness have resulted in changes in the function and structure of the eye in some astronauts which has been described as Spaceflight-Associated Neuro-ocular Syndrome (SANS). The leading hypothesis is that ocular changes associated with SANS result from chronic exposure to the weightlessness-induced headward fluid shift. The resulting venous congestion in the upper body may impair cerebrospinal and lymphatic fluid drainage from the skull and contribute to some of the changes in the eye. Further, venous stasis may develop in some astronauts, increasing the risk of a venous thrombus. Countermeasures that reverse the headward fluid shift during spaceflight such as centrifugation or lower body negative pressure have been proposed as means to relieve venous congestion associated with weightlessness and thus may mitigate the risk of SANS, venous thrombosis, and other spaceflight-induced cardiovascular adaptations. However, the amount of fluid shift reversal required to prevent SANS, venous thrombosis, and cardiovascular adaptations is unresolved. Parabolic flight provides a unique opportunity to evaluate the acute changes in the headward fluid shift within the venous compartment induced by varying levels of gravity during partial gravity and weightlessness.

The primary task objectives are to determine whether exposures to partial gravity levels (G-levels) similar to extraterrestrial levels (Moon and Mars) will provide protection against the headward fluids shifts that may be associated with the development of SANS, venous thromboembolism (VTE), and cardiovascular deconditioning. With the results from this partial gravity parabolic campaign, we will provide additional data that will contribute to comprehensive models of the relationship between G-levels and internal jugular vein (IJV) dimensions, pressure, and flow. Furthermore, investigating IJV hemodynamics bilaterally and studying the IJV in these subjects during parabolas producing weightlessness will better inform NASA regarding the risk of SANS and venous thrombosis and aid in the development of appropriate countermeasures.

Specific Aims

- 1. Determine the relationship between G-level and upper body venous hemodynamic parameters associated with a headward fluid shift, including IJV pressure, flow pattern, and cross-sectional area.
- 2. Determine if the relationship between G-level and upper body venous hemodynamic parameters is different between the left and right sides of the body.
- 3. Determine whether the supine posture in normal gravity produces similar internal jugular vein hemodynamics as occurs in acute weightlessness.

This research is directed because it contains highly constrained research. This work was recommended by the Human

Health Countermeasures (HHC) element to be performed as a directed study (a) to fulfill the partial gravity flight campaign human data collection objectives outlined in the previously-selected NASA Research Announcement (NRA; NNJ15ZSA001N-AG); (b) to provide the necessary data to accelerate HHC Spaceflight Associated Neuro-ocular Syndrome (SANS) risk reduction; and (c) to provide the necessary headward fluid shift data to help inform requirements for future lunar (Artemis Program) and Mars exploration. This proposed task will allow Human Research Program (HRP) to take advantage of a planned European Space Agency (ESA)-sponsored partial gravity parabolic flight opportunity in Spring or Summer of 2022. Furthermore, utilizing the same study design and outcome measures as the first campaign will ensure data consistency, enable us to leverage previously-developed human research documentation, hardware, and engineering analyses required by the parabolic flight provider (cost efficiency). When our first study was selected as part of a NASA Research Announcement (NRA) (NNJ15ZSA001N-AG), the power analysis indicated that data should be collected from 15 subjects. However, the ESA parabolic flight plan limited data collection to 9 subjects. Due to technical constraints and reduced operator support in the first campaign, internal jugular vein (IJV) area was obtained in only 8 of 9 subjects and IJV pressure only in 3 subjects at the lower G-levels. Thus, collecting data from 9 additional subjects as outlined here will allow us to fully explore the threshold G-level required to prevent venous congestion (IJV area and pressure) and altered flow patterns. In addition, this will allow us to assess the effects of right vs. left IJV anatomy on IJV dimension, pressure, and flow to varying G-levels, which has not been previously explored and may be key in assessing SANS and thrombosis risks. Finally, the previous partial gravity parabolic flight campaign did not include parabolas producing weightlessness, and thus the previous dose-response curves describing the relationship between G-level and IJV area required the assumption that IJV dimensions in the supine posture was a suitable analog. With the addition of weightlessness parabolas in a new campaign, this approach will be validated. Data from this project also will complement the weightless parabolic flight study recently funded for our laboratory (Characterization of Jugular Venous Blood Flow during Acute Fluid Shifts, Principal Investigator: Karina Marshall-Goebel, Ph.D., 80JSC019N0001-OMNIBUS - HERO Appendix B: NASA Human Research Program Omnibus Opportunity).

Rationale for HRP Directed Research:

**Task Description:** 

Research Impact/Earth Benefits:

The primary task objectives are to determine whether exposures to partial G-levels (similar to those on the Moon and Mars) will provide protection against the headward fluids shifts that may be associated with the development of SANS, venous thromboembolism, and cardiovascular deconditioning. With the results from this parabolic flight campaign, we will provide additional data that will contribute to comprehensive models of the relationship between G-levels and IJV dimensions, pressure, and flow and inform countermeasure development.

**Task Progress:** 

In this first year of the study, the team achieved several key milestones. First, the team submitted and received approval of the NASA Institutional Review Board documents, including the Informed Consent form. The consent forms to be reviewed by the French National Comité de Protection des Personnes (NCPP) will be completed in the Spring of 2023. Second, the team delivered the first draft of the Experiment Safety Data Package (ESDP), which included an overview of the study objectives and methods as well as preliminary diagrams depicting the workstations for the test subjects and sonographers. In consultation with Novespace (Bordeaux, France), the team redesigned the workstations to facilitate imaging of the internal jugular veins bilaterally while improving the ergonomics of probe placement and ultrasound keyboard manipulation. In the coming months, the team will finalize the workstation design, finalize the ESDP with Novespace, train the operators for data collection, and develop the data analysis plan to facilitate data reduction and reporting. Thereafter, the team will prepare hardware for shipment to and from Novespace, participate in the Flight Readiness Review, collect the data, and begin data reduction. Data analysis and reporting will be completed in the final year of this study.

The parabolic flight campaign is scheduled for June 2023.

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Bibliography Type:	Description: (Last Updated: 02/22/2024)
Abstracts for Journals and Proceedings	Lee SMC, Miller CA, Martin DS, Laurie SS, Lytle J, Young M, Macias BR. "Determining dose response profile of the headward fluid shift during varying gravity levels." NASA Human Research Program Investigators' Workshop, To the Moon: The Next Golden Age of Human Spaceflight, Galveston, Texas, February 7-9, 2023. NASA Human Research Program Investigators' Workshop, To the Moon: The Next Golden Age of Human Spaceflight, Galveston, Texas, February 7-9, 2023. , Jan-2023