X14 1 X7	EN/ 2021		TX 10/20/2020
Fiscal Year:	FY 2021	Task Last Updated:	FY 10/29/2020
PI Name:	Brainard, George C. Ph.D.		
Project Title:	Testing Solid State Lighting Countermeasures to Improve Circadian Adaptation, Sleep, and Performance During High Fidelity Analog and Flight Studies for the International Space Station		
Division Name:	Human Research		
Program/Discipline:			
Program/Discipline Element/Subdiscipline:	HUMAN RESEARCHBehavior and performance		
Joint Agency Name:		TechPort:	Yes
Human Research Program Elements:	(1) HFBP:Human Factors & Behavioral Performance	ce (IRP Rev H)	
Human Research Program Risks:	 (1) BMed:Risk of Adverse Cognitive or Behavioral Conditions and Psychiatric Disorders (2) Sleep:Risk of Performance Decrements and Adverse Health Outcomes Resulting from Sleep Loss, Circadian Desynchronization, and Work Overload 		
Space Biology Element:	None		
Space Biology Cross-Element Discipline:	None		
Space Biology Special Category:	None		
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Zip Code:	19107-5083	Congressional District:	1
Comments:			
Project Type:	FLIGHT	Solicitation / Funding Source:	2013-14 HERO NNJ13ZSA002N-BMED Behavioral Health & Performance
Start Date:	12/01/2014	End Date:	08/31/2022
No. of Post Docs:	0	No. of PhD Degrees:	
No. of PhD Candidates:	3	No. of Master' Degrees:	
No. of Master's Candidates:	1	No. of Bachelor's Degrees:	
No. of Bachelor's Candidates:	0	Monitoring Center:	NASA JSC
Contact Monitor:	Whitmire, Alexandra	Contact Phone:	
Contact Email:	alexandra.m.whitmire@nasa.gov		
Flight Program:	ISS		
	Flight Definition NOTE: End date changed to 8/31/2022 per NSSC in	nformation (Ed., 8/31/21)	
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Flight Assignment:	NOTE: End date changed to 03/31/2020 per NSSC information (Ed., 9/3/19)		
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	NOTE: End date changed to 11/30/2018 per NSSC		,
	NOTE: Element change to Human Factors & Behav (Ed., 1/17/17)	vioral Performance; previo	usly Behavioral Health & Performance

Key Personnel Changes/Previous PI:	December 2019 - Smith Johnston, MD, retired from NASA but has stayed active on this project.
COI Name (Institution):	 Barger, Laura Ph.D. (Brigham and Women's Hospital/Harvard Med Ctr) Clark, Toni B.S. (NASA Johnson Space Center) Czeisler, Charles M.D., Ph.D. (Brigham and Women's Hospital/Harvard Medical Center) Johnston, Smith M.D. (NASA Johnson Space Center (Retired 12/19, but still involved)) Moomaw, Ronald O.D. (NASA Johnson Space Center) Lockley, Steven Ph.D. (Co-PI: Brigham and Women's Hospital) Hanifin, John Ph.D. (Thomas Jefferson University) Rahman, Shadab Ph.D. (Brigham and Women's Hospital) St Hilaire, Melissa Ph.D. (Brigham and Women's Hospital)
Grant/Contract No.:	NNX15AC14G
Performance Goal No.:	
Performance Goal Text:	
Task Description:	This research addresses the NASA Research Announcement (NRA) NNJ13ZSA002N-BMED: Behavioral Health and Human Performance: "Evaluation of the Neurobehavioral Effects of a Dynamic Lighting System on the ISS." This NRA solicited both "Ground Based and Flight-Definition" research with the specific instructions that the "ground study serves as a precursor to the flight study, therefore the ground study solud take place in an analog with high fidelity to the ISS. The SSLAs should be studied in a high fidelity ground analog environment, then implemented on ISS to evaluate individual crewmember outcomes related to circadian physiology, sleep, behavioral health and performance using sensitive and validated measures that are feasible in the spaceflight environment." Currently, the International Space Station (ISS) uses General Luminaire Assemblies (GLAs) that house fluorescent lamps for illuminating the astronauts' working and living environments. NASA has determined that, beginning in 2016, the GLAs would be replaced with Solid-State Light Assemblies (SSLAs) containing Light Emitting Diodes (LEDs). Engineers at Kennedy Space Center developed a prototype Solid-State Lighting Assembly (SSLA) that was successfully installed onboard the ISS during ISS Expedition 18. The Principal Investigator and Co-Principal Investigator of the intender descarch worked with engineers, scientists, and mangers from Johnson Space Center (JSC) to revise the SSLA specifications so that the new lighting units would have dual capacity to: 1) provide illumination for crew members' working and living quarters, and 2) serve as a lighting countermeasure for crewmembers' circadian and sleep disruption. NASA ordered and received for a set of SSLAs intandrized psychometric, physiological, and neurobehavioral measures are testing the efficacy of light from the SSLA tak induction to the SSLA stake theore the state of the SSLA retrofit, the investigators have started the inflight ISS study to assess the acceptability use, and impact of deployment of adynami

Rationale for HRP Directed Research:

The sleep deficits experienced by astronauts during spaceflight along with risk of incompatible habitat design can be considered threats to the success of space missions (NASA Human Research Program Integrated Risk Plan, 2020). The resulting physiological and behavioral changes caused by sleep and circadian disruption can lead to diminished alertness, cognitive ability and psychomotor performance (Dijk et al., Amer. J. Physiol., 2001; Human Health and Performance Risks of Space Exploration Missions. McPhee and Charles, eds., 2010). As a measure to counteract sleep disruptions, crewmembers report using sleep promoting drugs: 71% on space shuttle flights and 75% during ISS expeditions (Barger et al., Lancet Neurology, 2014; Flynn-Evans et al., 2016). A significant portion of the global population suffers from chronic sleep loss and/or circadian-related disorders. Evidence for disease occurring due to a disruption of circadian homeostasis has mounted significantly in the past several years. In the United States, nearly 22 million Americans do shift work that interferes with a biologically healthy nocturnal sleep cycle (US Bureau of Labor Statistics, 2007). It has been shown that shift workers are more likely to suffer from a wide variety of ailments, including cardiovascular disease, metabolic disorders, gastrointestinal distress, and cognitive and emotional problems. Development of an in-flight lighting countermeasure that helps resolve circadian and sleep disruption in astronauts is likely to help optimize the use of light therapy for patient populations with affective, circadian and sleep disorders.

References

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McPhee J and Charles J, eds. 2010. Human Health and Performance Risks of Space Exploration Missions: Evidence Reviewed by the NASA Human Research Program. NASA SP-2009-3405 edition.

Barger LK, Flynn-Evans EE, Kubey A, Walsh L, Ronda JM, Wang W, Wright KP and Czeisler CA (2014) Prevalence of sleep deficiency and use of hypnotic drugs in astronauts before, during, and after spaceflight: an observational study. Lancet Neurology 13:904-912.

Flynn-Evans EE, Barger LK, Kubey AA, Sullivan JP and Czeisler CA (2016) Circadian misalignment affects sleep and medication use before and during spaceflight. npj Microgravity 2:15019; doi:15010.11038/npjmgrav.12015.15019.

Grant Establishment: Three institutions are collaborating on this multidisciplinary research: Thomas Jefferson University (TJU) in Philadelphia, Brigham and Women's Hospital (BWH) in Boston, and Johnson Space Center (JSC) in Houston. The start date for the grant was December 1, 2014. Subcontracts were then established between TJU, BWH, and Lockheed Martin. The aims were to complete a ground-based study in a high fidelity analog of the crew sleeping quarters (CQ), and an in-flight study in the daily living environment of the ISS.

Ground Based Analog Study: This study aims tested the efficacy of lighting protocols for daily operations using Solid State Lighting Assemblies (SSLAs) in ISS CQs installed in laboratories at TJU. In a controlled 5-day inpatient study using astronaut-aged volunteers, we tested the hypotheses that compared to the static, daily lighting of General Illumination only, the Dynamic Lighting Schedule protocol for a typical ISS work day (18 h wake: 6 h sleep) will improve visual performance, circadian entrainment, onset of melatonin production, sleep onset, sleep duration as well as morning alertness and performance. Separate human use protocols were submitted and approved by the Institutional Review Boards (IRBs) at TJU and NASA. Previously, NASA and National Space Biomedical Research Institute (NSBRI) funded the PI and Co-PI to develop a high fidelity, in-laboratory analog environment to study the visual, biological, and behavioral effects of the SSLAs. Specifically, a high-fidelity replica of the ISS Crew Sleeping Quarters (CQ) was developed with precise replication of CQ volume, geometry, and surface reflectance with an SSLA providing illumination. Astronaut-aged study subjects were able to be upright in this CQ and work, read, or use a computer just as crewmembers do onboard the ISS. In addition, a second CQ was developed that allows subjects to be semi-recumbent during wakefulness in SSLA lighting or fully recumbent when sleeping in darkness. Data from controlled studies in these high fidelity in-laboratory analog conditions represent the only published ground-based human data on the efficacy of the SSLAs to date (Brainard et al., Acta Astronaut., 2013; Brainard et al., Curr. Opin. Pulm. Med., 2016). In that earlier work, however, only a single subject could be studied at a time in the facility. In the current work, a second high-fidelity recumbent CQ was built and installed in the test facility enabling us to study up to two subjects at a time, with the intent of improving our speed for acquiring data in the analog facility. The SSLAs were each adjusted for their spectral output to be as close as possible to the NASA's vendor requirements for ISS (NASA Revision C, S684-13489, 2013). These specifications include Correlated Color Temperature (CCT or K) and luminance in candelas (cd) for three basic settings: 1) General Illumination: 4500 K SSLA white light, 210 cd; 2) Phase Shift/Alertness: 6500 K SSLA (blue-enriched) white light, 420 cd; 3) Pre-Sleep: 2700 K SSLA (blue-depleted) white light, 90 cd.

Based on published and unpublished data, the Co-PIs determined that the 90 cd luminance at crewmember's eye level inside of a CQ would be too bright to serve as an effective Pre-Sleep countermeasure. This issue was discussed with our project management team at JSC on several occasions. It was determined that in spaceflight, the SSLA luminance could be lowered from 90 cd using a combination of SSLA dimming buttons and a cloth shade system that is currently used on the fluorescent lighting system in the CQs onboard ISS. Based on a series of SSLA lighting measures and our prior pilot study in the CQs, we chose a Pre-Sleep luminance of 7.7 cd (20 lux at eye level) for our Pre-Sleep setting.

This study included male and female volunteers in good physical and mental health with normal color vision. Volunteers were selected in the age range of astronauts (range 26-54 years). Prior to admission to the laboratory, subjects were asked to maintain a regular 8:16 h, sleep:wake schedule and wear a wrist-borne, non-invasive activity and light monitor for at least 10 days. Over 550 subjects expressed interest in participating in this study. Approximately half of those individuals were not eligible based on phone interviews. Among those who were potentially eligible, 70 signed consent paperwork. Twenty-eight of those subjects completed the screening process and were randomly assigned into a lighting condition of either dynamic (N=16) or static (N=12) lighting. Among those subjects, 19 were male and 9 were female (age range 26 - 53 years). Twenty-five of these subjects successfully completed the entire five day study.

The data gathered from this first study run include successful collection of complete pre-study actigraphy, and inpatient study actigraphy from each subject. A total of 268 neurocognitive and performance tests were collected from each subject across the inpatient study (over 6,700 total). In addition, 95 Karolinska sleepiness scales (KSS) were collected from each subject across the inpatient study (2,375 total). Complete sets of blood, saliva, and urine samples were collected from each subject for the measurement of melatonin and 6-sulfatoxymelatonin. Melatonin contents of 548 plasma samples from 25 participants who completed a full study run have been analyzed. Polysomnography (PSG) was used to monitor sleep states and wakefulness using electrodes placed on the scalp, face, chin, and chest. Electrodes were positioned according to the International 10-20 System. The actigraphy, neurocognitive, and performance tests and urinary 6-sulfatoxymeltonin measures match similar or identical tests that were used onboard ISS during the flight study. Subject recruitment and enrolling is now completed. Data analysis is in process on the numerous dependent variables from the 5-day study. Portions of data from this study have been uploaded to the NASA Large File Transfer service.

Task Progress:

The testing of visual performance and color vision under different SSLA light settings has been done separately from the five day studies. Two separate cohorts of 8 healthy male and female, astronaut-aged subjects have completed within-subjects study designs that test their visual performance and color vision. The data from each of these studies is being analyzed.

ISS Flight Study: The aims of this study are to test the efficacy of lighting protocols for daily operations using SSLAs for inflight crewmembers onboard ISS missions. Specifically, we are assessing the acceptability, use and operational impact of deployment of the Dynamic Lighting Schedule protocol on astronaut vision, sleep, alertness, circadian

	rhythms, and general well-being during ISS flight missions. This inflight study will test the hypotheses that, compared to current static daily fluorescent lighting of General Illumination only, the Dynamic Lighting Schedule protocol will maintain acceptable visual performance and color discrimination for operational tasks, improve circadian entrainment, improve circadian adaptation following a sleep shift challenge such as a 'slam-shift,' improve sleep duration and efficiency, and enhance wake-time alertness and cognitive performance.
	Ethical approvals were obtained from NASA and Partners Healthcare for the flight study. The flight study successfully went through an ISS Medical Project (ISSMP) feasibility assessment. Subsequently, the Human Research Program (HRP) Science Management Panel selected this study for flight on 9/3/15. The first ISSMP teleconference was held on 9/30/15 involving representatives from JSC's ISSMP, BWH, and TJU. This teleconference was held monthly through the set up and running of the study. A total of eight crew members consented to participate in the flight study. Seven astronauts have completed pre-flight, in-flight, and post-flight testing. As of October, 2020, 77 SSLAs have been successfully installed providing a 92% retrofit of the ISS.
	This ISS flight study on crewmembers is a sophisticated human photobiological study. All photobiological studies, whether in spaceflight or on Earth, rely on precise characterization of the independent variable of the study: light. For this study, the relevant light stimulus is light emitted by the new SSLAs and the remains of the original ISS fluorescent lighting system. A spectrophotometer/irradiance meter is an essential tool for ensuring that consistent emission of light spectrum and light intensity are maintained during the inflight ISS research. The key measures for this flight study are light irradiance, and spectral power distribution of the four settings of the SSLAs, as well as the single setting of the current fluorescent lights. Working with the study collaborators, ISSMP has selected and deployed the meter for the flight study on ISS. A total of 37 sets of lighting measures have been taken by crewmembers and transmitted from ISS to the study team on Earth.
	Considerable work between the study collaborators and the hardware group of Research Operations & Integration (ROI) element [Ed. note October 2020: ROI was formerly ISSMP; element change in 2019) has gone into making flight worthy versions of visual performance and color vision tests. Those tests were used successfully on the ISS. The Lanthony Desaturated 15-Hue test is the method to be used for testing crewmember color discrimination under the different SSLA light settings compared to that of the current fluorescent lighting on ISS. Historical data of actigraphy, sleep logs, cognitive testing, and urine samples have been identified from previous flight studies that will act as the control data in the flight study. Discussions are ongoing about data sharing for these historical data. Through September 2019, seven crewmembers have completed all post-flight data collection, including actiwatch/sleep logs, cognition, urine sampling, and visual testing. Data are being analyzed on these dependent measures. The urine samples have been returned to Earth and will be analyzed for 6-sulfatoxymelatonin at BWH in the near range. After completion of their flight mission, each of the seven crewmembers met personally with one or two representatives from BWH and TJU for debriefing at JSC. The project is scheduled to end on December 31, 2020. It is anticipated that data analysis and development of manuscripts will continue after that end date.
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Bibliography Type:	Description: (Last Updated: 10/30/2023)
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Abstracts for Journals and Proceedings	Kent BA, Rahman SA, St Hilaire MA, Clark T, Hanifin JP, Barger LK, Czeisler CA, Brainard GC, Lockley SW. "The effects of dynamic lighting on urinary 6-sulphatoxymelatonin rhythms during the NASA HERA campaign." Harvard University, Division of Sleep Medicine, Farrell Prize Day, October 28, 2019. Harvard University Farrell Prize Day, October 28, 2019. , Oct-2019
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Articles in Peer-reviewed Journals	Grant LK, Gooley JJ, St Hilaire MA, Rajaratnam SMW, Brainard GC, Czeisler CA, Lockley SW, Rahman SA. "Menstrual phase-dependent differences in neurobehavioral performance: the role of temperature and the progesterone/estradiol ratio." Sleep. 2020 Feb 13;43(2):zsz227. <u>https://doi.org/10.1093/sleep/zsz227</u> ; <u>PMID: 31670824</u> ; <u>PMCID: PMC7457328</u> , Feb-2020
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Awards	Brainard G. (George Brainard, PhD) "Fellow of the Illuminating Engineering Society (IES) on August 9, 2019 in Louisville, KY. This honor is a lifetime award that formally recognizes his valuable contributions to the technical activities of the IES and to the broader lighting industry." Aug-2019
Books/Book Chapters	Barger LK, Dinges DF, Czeisler CA. "Sleep and Circadian Effects of Space." in "Handbook of Bioastronautics." Ed. L.R. Young, J.P. Sutton. Cham: Springer, first online 24 October 2020. https://doi.org/10.1007/978-3-319-10152-1_86-2, Oct-2020