

<b>Fiscal Year:</b>	FY 2016	<b>Task Last Updated:</b>	FY 10/06/2016
<b>PI Name:</b>	Fuller, Charles A. Ph.D.		
<b>Project Title:</b>	Head-Down Tilt as a Model for Intracranial and Intraocular Pressures, and Retinal Changes during Spaceflight		
<b>Division Name:</b>	Human Research		
<b>Program/Discipline:</b>	HUMAN RESEARCH		
<b>Program/Discipline--Element/Subdiscipline:</b>	HUMAN RESEARCH--Biomedical countermeasures		
<b>Joint Agency Name:</b>	<b>TechPort:</b>	No	
<b>Human Research Program Elements:</b>	(1) <b>HHC</b> :Human Health Countermeasures		
<b>Human Research Program Risks:</b>	(1) <b>SANS</b> :Risk of Spaceflight Associated Neuro-ocular Syndrome (IRP Rev I)		
<b>Space Biology Element:</b>	None		
<b>Space Biology Cross-Element Discipline:</b>	None		
<b>Space Biology Special Category:</b>	None		
<b>PI Email:</b>	<a href="mailto:cafuller@ucdavis.edu">cafuller@ucdavis.edu</a>	<b>Fax:</b>	FY
<b>PI Organization Type:</b>	UNIVERSITY	<b>Phone:</b>	530-752-2979
<b>Organization Name:</b>	University of California, Davis		
<b>PI Address 1:</b>	Section of Neurobiology, Physiology & Behavior		
<b>PI Address 2:</b>	One Shields Avenue, 196 Briggs Hall		
<b>PI Web Page:</b>			
<b>City:</b>	Davis	<b>State:</b>	CA
<b>Zip Code:</b>	95616-5270	<b>Congressional District:</b>	3
<b>Comments:</b>			
<b>Project Type:</b>	GROUND	<b>Solicitation / Funding Source:</b>	2011 Crew Health NNX11ZSA002NA
<b>Start Date:</b>	02/01/2013	<b>End Date:</b>	12/31/2017
<b>No. of Post Docs:</b>	0	<b>No. of PhD Degrees:</b>	2
<b>No. of PhD Candidates:</b>	0	<b>No. of Master' Degrees:</b>	0
<b>No. of Master's Candidates:</b>	0	<b>No. of Bachelor's Degrees:</b>	0
<b>No. of Bachelor's Candidates:</b>	0	<b>Monitoring Center:</b>	NASA JSC
<b>Contact Monitor:</b>	Allcorn, Aaron	<b>Contact Phone:</b>	281.244.8402
<b>Contact Email:</b>	<a href="mailto:aaron.j.allcorn@nasa.gov">aaron.j.allcorn@nasa.gov</a>		
<b>Flight Program:</b>			
<b>Flight Assignment:</b>	NOTE: End date changed to 12/31/2017 per NSSC information (Ed., 4/20/2016) NOTE: End date will be 6/30/2017 per R. Brady/JSC HRP (Ed., 11/3/15)		
<b>Key Personnel Changes/Previous PI:</b>			
<b>COI Name (Institution):</b>	Hoban-Higgins, Tana ( University of California, Davis ) Murphy, Christopher ( University of California, Davis ) Robinson, Edward ( University of California, Davis ) Gompf, Heinrich ( University of California Davis )		
<b>Grant/Contract No.:</b>	NNX13AD94G		
<b>Performance Goal No.:</b>			
<b>Performance Goal Text:</b>			

<b>Task Description:</b>	<p>This ground-based program is intended to address the etiology of visual system structural and functional changes observed in astronauts during both inflight and postflight periods. Using the well-documented rat hindlimb suspension (HLS) model, functionally equivalent to human head-down bedrest, we will examine the relationship between cephalic fluid shifts resulting from long-duration G-unloading and the regulation of intracranial and intraocular pressures, as well as the effects these same cephalic fluid shifts have on visual system structure and function. Animals will be chronically instrumented with biotelemetry to continuously measure intracranial pressure. Additionally, regular intraocular pressure measurements will be made by tonometry during long-term exposure to cephalic fluid shifts induced by suspension. MRI images visualizing the visual system morphology will also be collected from HLS and control animals at regular intervals. Retinal morphology and ultrastructure will be examined at specified intervals both during HLS and post-HLS recovery by both ophthalmic examinations and tissue histology evaluation. Changes in retinal/visual function will be regularly assessed electrophysiologically by measuring visual evoked potentials and electroretinograms. This program will utilize both male and female subjects in order to examine possible gender differences in these responses. We will also examine the possible contributory factors of aging and elevated atmospheric carbon dioxide (hypercapnia) on to these responses of the visual system. Further, in addition to mimicking the effects of long duration exposure to microgravity through the use of the HLS model, we will examine the responses of our measured outcomes during long-term recovery in the post-HLS period. Collectively, these data will help allow us to develop a model to both understand and predict the etiology of changes in visual structure and function in astronauts exposed to the microgravity of spaceflight and during postflight recovery. In summary, our ultimate goal is to develop a translational mammalian model by which the data generated using this model can facilitate the development of countermeasures to alleviate any visual system decrements arising from exposure to the microgravity spaceflight environment.</p>
<b>Rationale for HRP Directed Research:</b>	
<b>Research Impact/Earth Benefits:</b>	<p>This research has the potential to help further our understanding of chronic cephalic fluid shifts on neurological and ophthalmic health. No innovative technologies have been developed during this period.</p>
<b>Task Progress:</b>	<p>This program utilizes both male and female subjects in order to examine possible gender differences in these responses. We are examining the possible contributory factors of aging and elevated atmospheric carbon dioxide (hypercapnia) on to these responses of the visual system. Further, in addition to mimicking the effects of long duration exposure to microgravity through the use of the HLS model, we are examining the responses of our measured outcomes during long-term recovery in the post-HLS period. Collectively, these data will help allow us to develop a model to both understand and predict the etiology of changes in visual structure and function in astronauts exposed to the microgravity of spaceflight and during postflight recovery. In summary, our ultimate goal is the development of a translational mammalian model; the data generated using this model would be used to facilitate the development of countermeasures to alleviate any visual system decrements arising from exposure to the microgravity spaceflight environment.</p> <p>During this period of performance, the research team has been hired and trained, major equipment purchased, and facilities set-up. We have put into place the core techniques and capabilities that will be necessary for the successful execution and completion of this program. This has included establishing a flow for the employment of the tail suspension model using the pigmented Long-Evans rat. While our initial subjects were young males, we are also studying cohorts of young females (to examine possible gender differences) and older males (to examine possible age effects). The cohort that presents the most significant response to HLS will be studied in a hypercapnic environment (similar to that experienced on the International Space Station (ISS)). This will allow us to determine if there is a role of increased CO<sub>2</sub> exposure in the etiology of these visual changes. The subjects currently proposed for use in the hypercapnic study are older males.</p> <p>We are utilizing biotelemetry to record intracranial pressure. We have extensively revised the data acquisition software, which has been necessary to both improve the ease and accuracy of data collection, as well as to tailor the system to meet the needs of this research program. The first cohort from which the biotelemetry data were collected were older males.</p> <p>Additional measurements of visual system function include complete ophthalmic clinical exams, measurement of intraocular pressure by tonometry, and, ultimately, tissue histology. The retinal imaging performed during this program include both fundus imaging with fluorescein angiography and OCT.</p>
<b>Bibliography Type:</b>	Description: (Last Updated: 12/07/2018)