





Success Looks Like Us

MINDS 2023 Talk

Cellas Ari'ka Hayes, PhD

Postdoctoral Research Fellow Propel Scholar

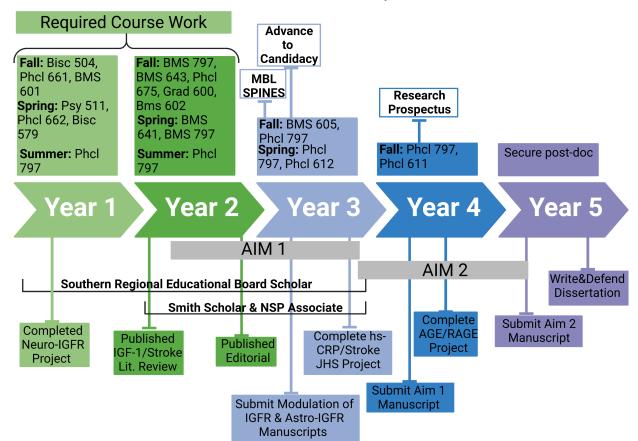
Department of Neurology and Neurological Sciences
School of Medicine
Stanford University

Life goes on but the scars remain – Lil Wayne

- Ludlow, MS
- Farm/rural life
- Single parent household
- Small K-12 school
- Gifted program
- Psychosis event
- Undergrad @ UofM
- First-Gen college graduate
- Graduate school @ UofM
- Finished in 3.5 as the 1st Black Male in History
- The Fam

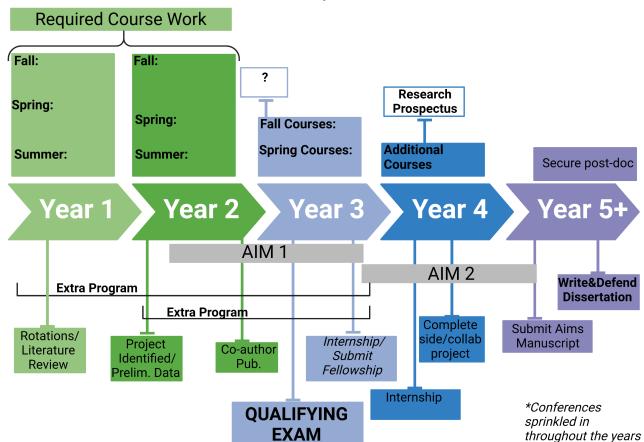


The Graduate School Snap Shot



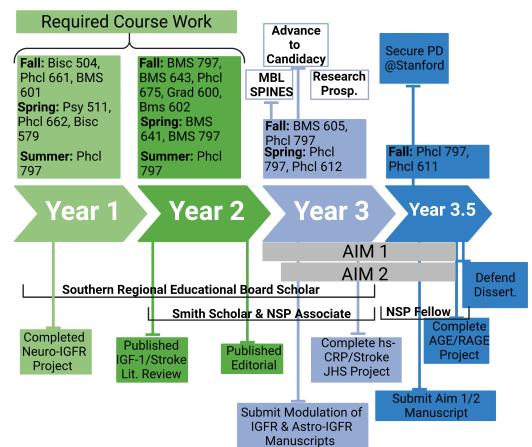


The Grad School Template



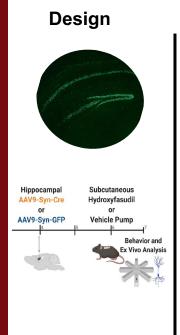


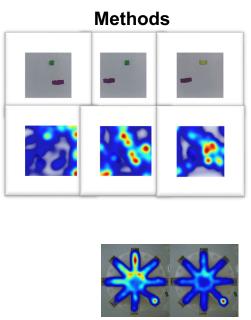
So... what did it actually look like for me?

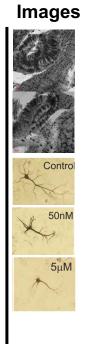




Fall 2016 – Spring 2019 Ashpole Lab University of Mississippi







Research Findings

- Male Cognitive Impairments in RAWM
- IGFR inhibition reduces neurite outgrowth
- IGFR inhibition alters Rho-Kinase Pathway

The Start of Belief

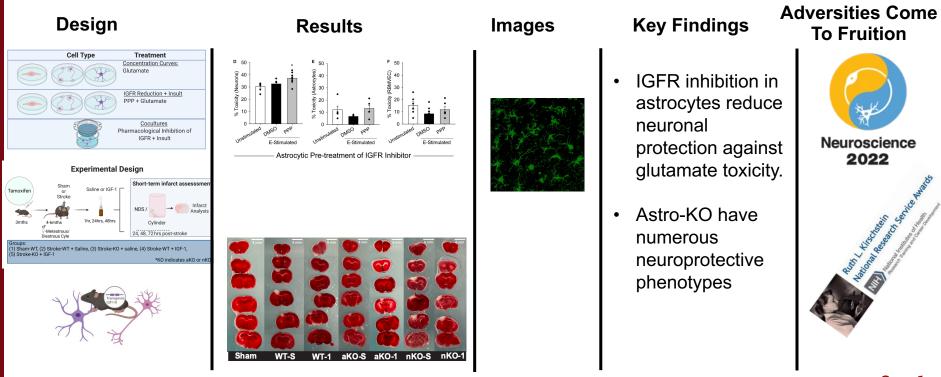








Graduate Research ~ University of Mississippi Ashpole Lab (2019-2022)



Graduate School Expectations? Is your neuroscience fulfilling? Navigating Challenges?



You Learn More In Failure Than You Ever Do In Success – Jay Z

Successful Grants

- UM Grant
- NIH NRSA F31

Failed Grants

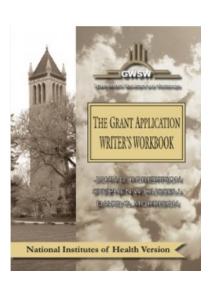
- NSF GRFP
- NSF GRFP
- Ford Predoctoral Fellowship
- American Heart Association Predoctoral Fellowship
- NIH NRSA F31

The Key: Grant and manuscript writing is code cracking. Once you get it, then you get it.



2019 Strength: Science Communication 2019 Weakness: Scientific Writing

- Scientific reading takes more strategy than it does skill.
- To develop a strategy, you need practice and time.



DISCUSSION

big summary/ mportance statement for the science

> scientific findings as extension of previous work

findings' scientific

how this engineering will facilitate future

forward-looking state-

engineering aspect of the

These results show that the capacity to induce Trees and modify their phenotype is a characteristic of more effector strains than was appre-ciated previously. Our findings concerning the role of human gut bacteria in shaping features of the gut mucosal immune system complement and extend the elegant work by Atarashi et al. (16). They used a single selective condition (chloroform treatment) to recover a group of 17 strains (all of which were described as members of the class Clostridia) from the human fecal microbiota of a single donor and showed that the consortium was capable of expanding the colonic regulatory T cell compartment in gnotobiotic mice. The fact that we found this effector activity among gut species belonging to other bacterial phyla suggests that distribution of this functional capacity may be beneficial in ensuring that this tolerogenic cell type is consistently

and persistently maintained in different microbial community and

host contexts. The approach we describe allows systematic follow-up

analyses of the extent to which the Tree response is affected by factors such as age at colonization or by different diets that produce abrupt and substantial alterations in microbiota configurations (45-47). Despite identifying members of different human gut bacterial phyla that shape the Tree response, our study and that of Atarashi et al. revealed that intestinal short-chain fatty acid concentrations increased upon colonization. Given the substantial amount of data supporting a role for short-chain fatty acids in the induction of Trees (42-44), this suggests a common pathway by which different microbes converge to modulate this facet of the host immune system. The genetic manipulability of some of the bacterial strains identified here, notably the Bacteroides, affords an opportunity to test this and other hypotheses, and advance our knowledge about the molecular underpinnings of

As the field of human microbial ecology research moves from

observational studies to hypothesis-driven experiments designed to

ments about the field as directly test the contributions of the microbiota and its components to health, there is a growing need to develop and transition to a mod-

ernized set of Koch's postulates (48) where the groups of microbes that modulate host phenotypic responses are identified along with the environmental factors (for example, dietary) necessary for the response to be fully manifest. We have developed a platform for systematically identifying microbe-host phenotype interactions in different (human) donor microbiota using gnotobiotic mice that can represent importance of the different host genetic features and different environmental conditions of interest. With the 17 strains in our culture collection, there were more than 100,000 possible combinations to search for effector strains. Using the mathematical and experimental strategies described, we only needed 100 combinations to identify multiple effector microbes for

microbiota-Tree crosstalk.

three very diverse biological responses (metabolic, adiposity, and Trees) This represents a 1000-fold reduction in the search space compared to what would be required theoretically. By testing these 100 combinations of microbes in an out-of-the-isolator gnotobiotic caging system rather than in traditional flexible film isolators, we overcame what would have been an insurmountable practical barrier to performing these studies for most groups. Our entire study could have been completed with a single flexible film isolator to generate the required germ-free mice. This feature suggests that our overall approach should be accessible to many investigators because animal facilities with small numbers of gnotobiotic isolators already exist in numerous universities.

Although identifying effector strains represents a critical first step

in mechanistic analyses of how the gut microbiota affects various

gnotobiotic mice genetically deficient in various components of the

immune system (such as Toll-like receptors or inflammasomes) and

facets of host biology, once such strains are identified, much additional work needs to be done. For example, numerous other important components of the intestinal immune system may also be affected by colonization with the strains we identified, including B cell class switching to IgA, macrophage/dendritic cell effector or migratory properties, and γδ T cell function. Another important goal is to identify the limitations of the effector molecules produced by the identified effector strains and the platform show which host signaling pathways through which these molecules act. Using questions would be

effector strains that are genetically manipulated (for example, through whole-genome transposon mutagenesis) represent ways for pursuing this goal. Although additional elements of these mechanistic analyses will be dependent on the biological processes being interrogated, in principle this platform can be applied to any microbiota-associated phenotype. Finally, our approach has therapeutic implications because it represents an enabling system for identifying and characterizing how it will facilinext-generation probiotics or combinations of pre- and probiotics

tate medical research

how it facili-

tates scientific

research

Faith et al., Science Translational Medicine (2014) doi:10.1126/scitranslmed.3008051

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Then & Now

Paper Tracking

Time Started Reading:	Time Finished Reading:	
Article Analysis-Title of Article:		
Abstract summary:		
Hypothesis of article:		
Figures analysis:		
Summary of Article and Analysis o	f Results:	

Thomas 2021

Title: Thomas 2021 Date: 2022-12-28 Time: 18:13 NEUROPATH

Association of TDP-43 proteinopathy, cerebral amyloid angiopathy, and Lewy bodies with cognitive impairment individuals with or without Alzheimer's disease neuropathology

Authors:

David X. Thomas 1,3*, Sumali Bajaj2,3, Kevin McRae-McKee2 , Christoforos Hadjichrysanthou/ M.Anderson2 & John Collinge 1

Year: 2021

tags/ key terms

#tdp43 #caa #lbd #AD

Introduction

Objective: Compare the association of common neuropathologies with pre-mortem cognitiv decline in the presence vs absence of concomitant Alzheimer's disease.

Methods

- NACC 2005-2018 #NACC #np
- CDR-SOB & MMSF
- Bayesian Hierarchical regression models to estimate the association of Alzheimer's diser neuropathology, TDP-43 proteinopathy, cerebral amyloid angiopathy (CAA), and Lewy be with cognitive trajectories afer accounting for the covariates consisting of demographic features, and other neuropathologies
- SIMILIAR VARIABLES AS CELLAS MANUSCRIPT PLAN AND DATA PROCESSING
- Binary categories were created for Alzheimer's disease neuropathological change, Lewy bodies, CAA, and TDP-43 proteinopathy using cut-of values which led to reasonably balanced groups.

Statistics

#bayesianhierarchicalregression

linear mixed effect models in a Bayesian framework

#linearmixedeffectregression #chisquare

Results

In ADNC+individuals, we assessed if the rate of cognitive decline was the same in the presence absence of a co-pathology, using one model each for TDP-43, CAA and Lewy Bodies. We estimated the two way interactions of TDP-43 proteinopathy and time (β MMSE=-0.34, 95% BCI (-0.64, -0.04); β CDR-SB =0.03, 95% BCI (0.13, 1.47)), CAA and time (β MMSE=-0.04, 95% BCI (-0.42, 0.35); β CDR-SB =0.09, 95% BCI (-0.11, 0.29)) and Lewy bodies and time (β MMSE=-0.25 95% BCI (-0.57, -0.01); β CDR-SB =0.19, 95% BCI (-0.01, 0.39)) (Fig. 2). These results suggest the ADNC+individuals with these pathologies have a steeper rate of cognitive decline compared to those without co-morbid pathologies.

However, our data showed no association of Lewy bodies with cognitive impairment over time ADNC- individuals.

Main Findings

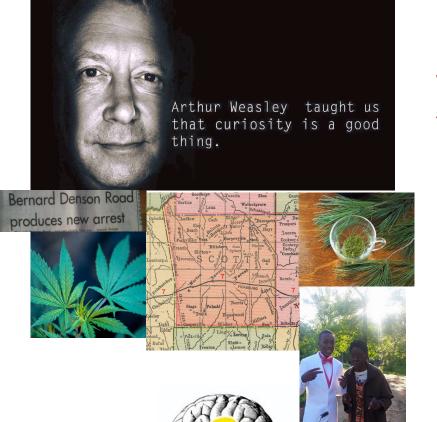
ADNC+individuals with these TDP-43, CAA, and lewy bodies individually have a steeper rate of cognitive decline compared to those without co-morbid pathologies. No association of Lewy bodies with cognitive impairment over time in ADNC- individuals.

IDEAS

non-binary dichotomizing variables to capture complexity of neuropathologies.

PDF





Where does your curiosity stem from?

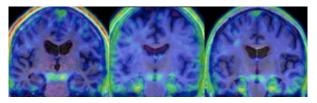
A Series of Unfortunate and/or Fortunate Events



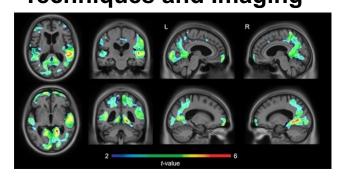
Why do you do what you do?

Postdoc Work- AD & Neurodegeneration & Vasculature & Neuroimaging

Research Interests



Techniques and Imaging



Support and Recruitment

Postdoctoral Recruitment Initiative in Science and Medicine

Stanford PRISM

Funding

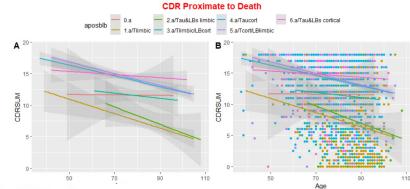
Propel,
Postdoctoral Scholars Program
https://propel.stanford.edu/

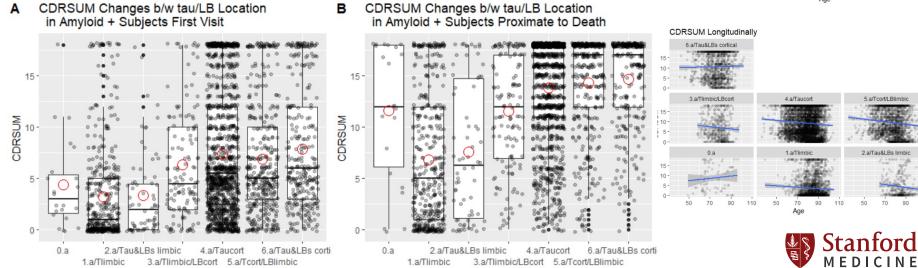


Postdoc Research

Pathology Progression Combination

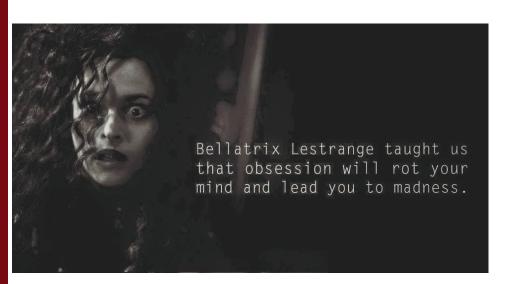
 Utilize NACC data to understand neuropathology of aging/AD





Pathology Progression Combination

Greatest lesson was time management and self-preservation through self-investment



- Pre-pandemic work hours
- Pandemic work hours
- Post-pandemic work hours
- Graduate school hours
- Postdoc hours

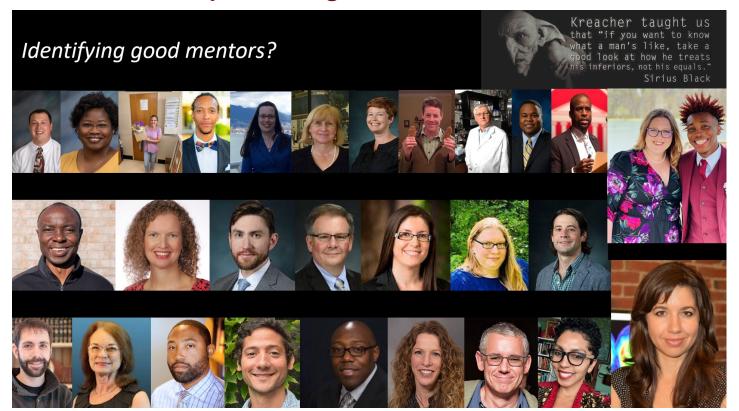


My Rewards versus Pitfalls of Academia

- 3 1st author publications
- 1st black trainee and 3rd in history (37 years)
- 5 1st coauthor manuscripts under review/inprep
- 25 honors, award, and features
- 9 noteworthy features, videos, sci-comm opportunities
- 6 oral presentations/ 6 poster presentations
- 120+ hours of community service
- 27 RCR hours
- 4 teaching hours
- 3 professional appointments
- 13 mentees

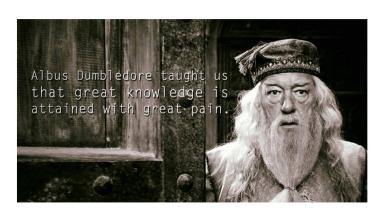


You are who you hang around -Whoever





"Obstacles" = "Opportunities"





Reading, Consolidating, Asking Questions, Opportunities

BMBH Emerging Scholars Program

















AAAS Science & Technology Policy Fellowships









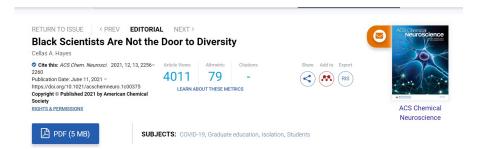
M.D. vs. Ph.D.

M.D.

- MCAT
- Years of courses where grades truly define you
- No room for extracurriculars
- Few research opportunities
- STEPS after STEPS
- Matching
- **\$\$\$**
- No room for intellectual development
- Few skills gained
- Stringent timeline
- Limited post-graduate opportunities



If you do choose a Ph.D., BE PREPARED



- THE UPROAR OF 2020, AN INTRODUCTION
 - A GRADUATE STUDENT PERSPECTIVE
- THE ADDITIONS OF BEING A MINORITY GRADUATE STUDENT
 - ARE BLACK STUDENTS THE BENEFICIARIES TO DEI WORK?
- THE FEELING OF OBLIGATION AND PROVING SELF-WORTH
 - BEING A SCIENTIST AND BEING A "BLACK SCIENTIST"
- BLACKNESS AS A STEPPING STONE AND THE REAL-WORLD PERSPECTIVE
 - BLACK STORIES AS TOKENS AND SPOTLIGHTS
- MAKING THE SYSTEM BETTER: A CONCLUSION AND GUIDE FOR THE FUTURE







A guide for applications

CELLAS HAYES. PHD

Be Genuine. Be Relatable. Build Your Life on Being Different.



CV AND APPLICATIONS AND PORTFOLIOS

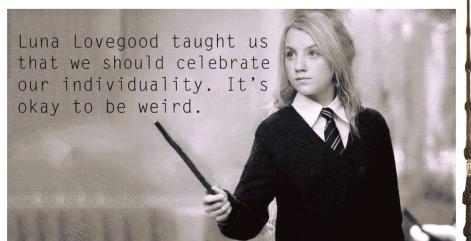
https://www.cellashayes.com/cv-and-applications



https://ars.sf.ucdavis.edu/sites/g/files/dgvnsk2741/files/inline-files/pre-grad-guide_1.pdf











Be Genuine Be Relatable Build Your Life on Being Different

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