

APDA'S RESEARCH AGENDA AND ACCOMPLISHMENTS

REBECCA GILBERT, MD, PHD

Chief Scientific Officer,
American Parkinson Disease Association

WHO WE ARE AND WHAT WE DO



**AMERICAN
PARKINSON DISEASE
ASSOCIATION**

Strength in optimism. Hope in progress.

AMERICAN PARKINSON DISEASE ASSOCIATION

Every day, we provide the support, education, and research that will help everyone impacted by Parkinson's disease live life to the fullest.

AMERICAN PARKINSON DISEASE ASSOCIATION

Local Programs & Services

- **Information and Referral**

- I&R Centers
- Toll free helpline 800-223-2732
- Ask the Doctor

- **Education & Support**

- Support groups
- Symposiums, Educational Events, Webinars

- **Health & Wellness**

- Exercise and movement classes
 - Dance, yoga, tai chi, boxing and more
- Fitness Professionals Training Program
- Creative arts programming

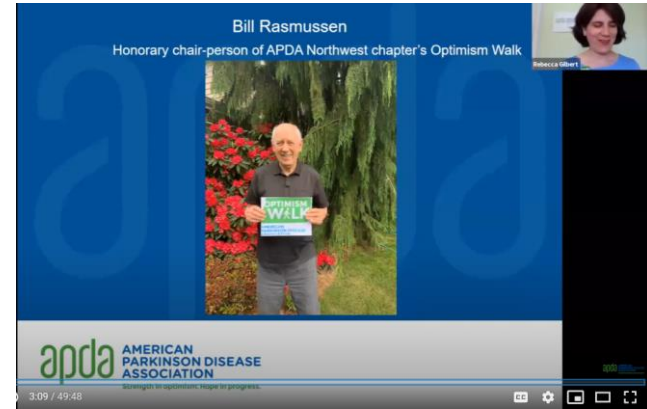


AMERICAN PARKINSON DISEASE ASSOCIATION

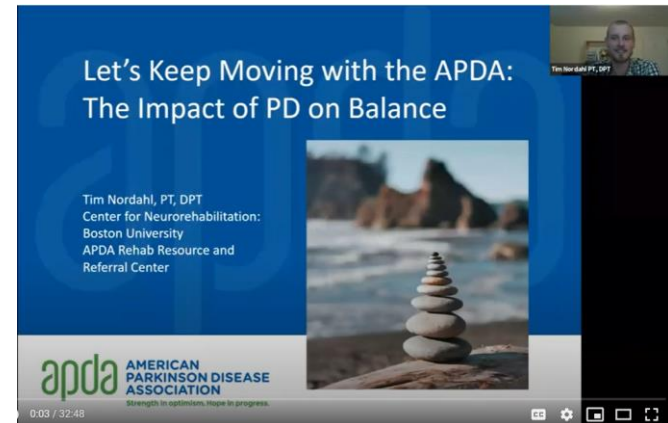
Online programming: education, support and wellness



Tai Chi with Craig (Level 2) 8/28/20



Dr. Gilbert Hosts



AMERICAN PARKINSON DISEASE ASSOCIATION

Nationwide Programs & Services

- Online resources
- Publications
- Webinars



Medications Approved for the Treatment of Parkinson's Disease in the USA

Below is a full list of Parkinson's medications that have been approved to treat Parkinson's in the United States, as of May 2018.

QUESTION & ANSWER



Un Jung Kang, MD
*Founders Professor of Neurology
Director of Translational Research*
The Marlene and Paolo Fresco Institute for
Parkinson's and Movement Disorders
NYU Langone Health
New York, NY



Cómo vivir bien con la enfermedad de Párkinson ¡10 cosas que puede hacer ahora!

Le han diagnosticado con la enfermedad de Párkinson. ¿Qué hacer ahora?

La enfermedad de Párkinson (EP) es un trastorno neurológico progresivo, sin embargo, la mayoría de las personas que tienen EP pueden vivir bien por muchos años con un buen plan de cuidado de la salud. Aunque no se ha comprobado si alguna terapia tiene un efecto "neuroprotector" o "modificador de la enfermedad", hay claros indicios de que la personas con EP pueden mejorar su calidad de vida si toman de inmediato medidas para fortalecer sus cuerpos y sus mentes. Mantenga una actitud positiva y comience desde ahora a poner en práctica las siguientes 10 medidas:

AMERICAN PARKINSON DISEASE ASSOCIATION - RESEARCH



George C. Cotzias Fellowship

Fellowship over three
years.



Post-Doctoral Fellowship

Fellowship for one
year.



Research Grants

Renewable one year
grant.



Diversity in PD Research Grant

Renewable one year
grant.

CENTERS FOR ADVANCED RESEARCH

APDA's Centers for Advanced Research support research trainees, fellowship programs, early-stage discovery and later-stage clinical translation. The Centers facilitate investigative research into the causes, treatments, and ultimately a cure for PD. The directors of each APDA Center (shown here) are among the most renowned in their field.

**Boston University
School of Medicine**
– Boston, MA
*Marie Hélène Saint-Hilaire,
MD, FRCP (C)*



**Brigham and Women's
Hospital**
– Boston, MA
Clemens Scherzer, MD



**University of Alabama
at Birmingham School
of Medicine**
– Birmingham, AL
David G. Standaert, MD, PhD



Mayo Clinic
– Jacksonville, FL
Dennis Dickson, MD



**Emory University
School of Medicine**
– Atlanta, GA
Thomas Wichmann, MD



**University of Pittsburgh
Medical Center**
– Pittsburgh, PA
*J. Timothy Greenamyre,
MD, PhD*



**Rutgers Robert Wood
Johnson School of
Medicine**
– New Brunswick, NJ
Mary Maral Mouradian, MD



**Washington University
School of Medicine**
– St. Louis, MO
Joel S. Perlmutter, MD

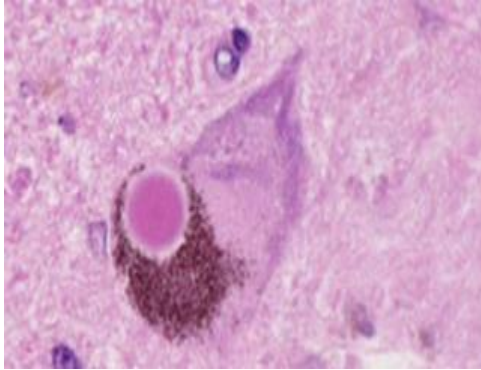


PD RESEARCH – PRECLINICAL AND CLINICAL

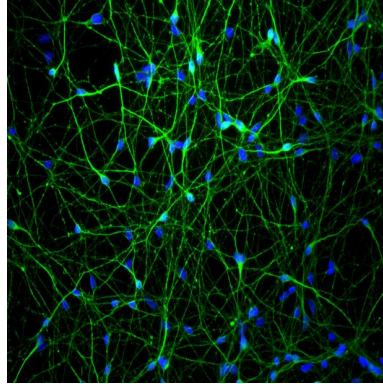
- Pre-clinical research - research involving cells, animal models, etc.
- Clinical research - research involving people



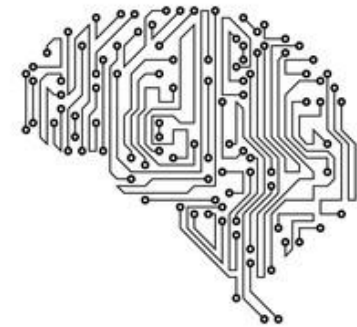
PARKINSON'S DISEASE: EXCITING THEMES IN CURRENT RESEARCH



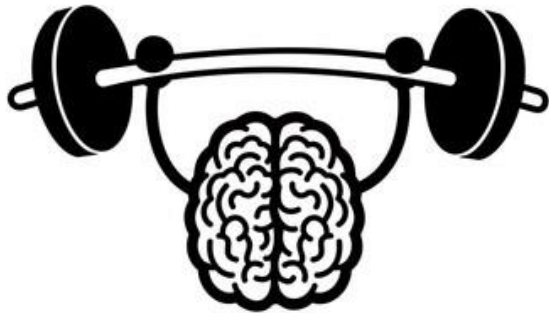
Abnormal accumulation of
alpha-synuclein



The role of other cell types
(not neurons)



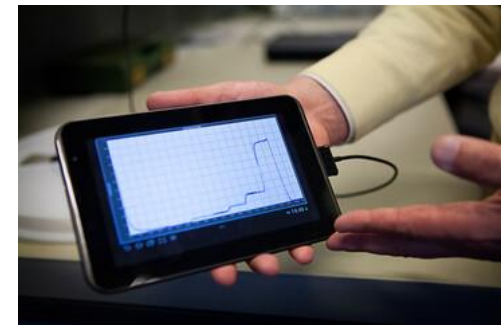
Brain circuitry



The role of exercise

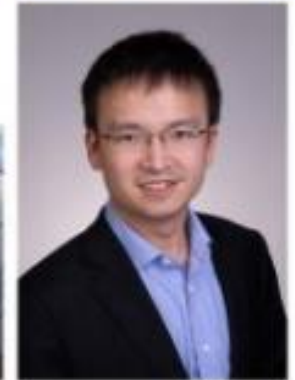


Genetics

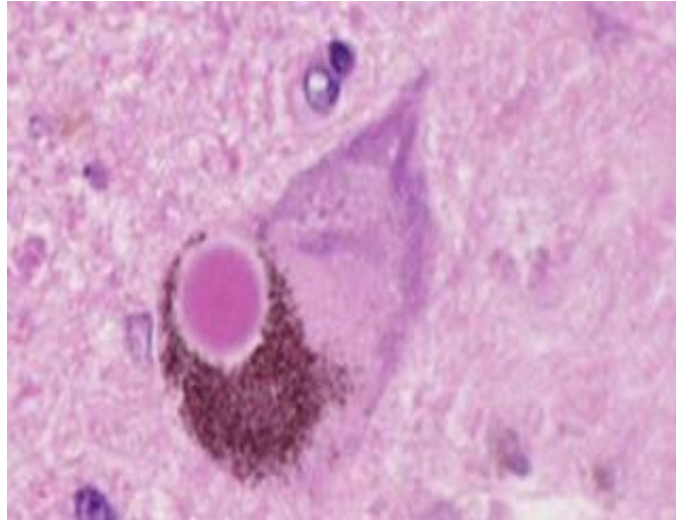


Digital biomarkers

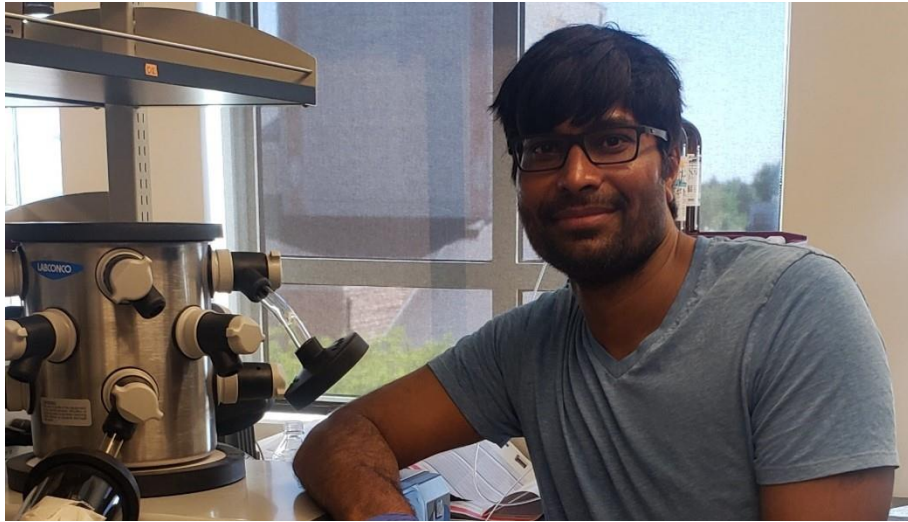
AMERICAN PARKINSON DISEASE ASSOCIATION - RESEARCH 2021



PARKINSON'S DISEASE AND ALPHA-SYNUCLEIN



HOW CAN WE STOP ALPHA-SYNUCLEIN FROM ACCUMULATING IN PD?

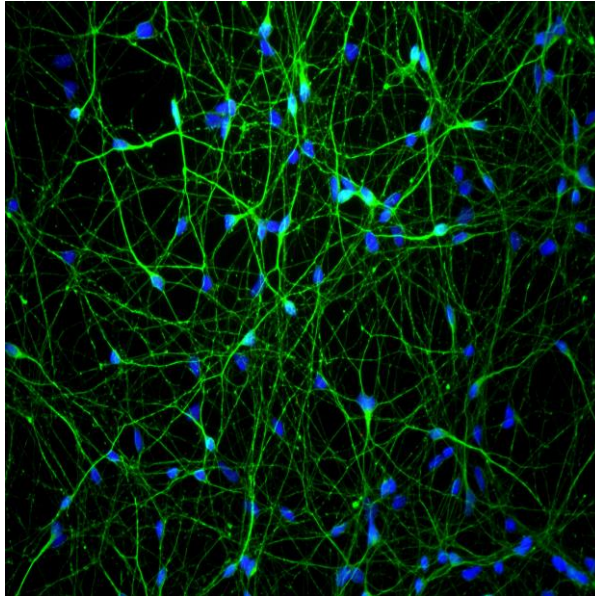


Dr. Sunil Kumar, University of Denver, Denver, CO

Objective:

To identify the portion of alpha-Synuclein that is essential for abnormal accumulation. The ultimate goal is to develop therapies that block this portion and prevent accumulation.

PARKINSON'S DISEASE AND OTHER BRAIN CELLS



PARKINSON'S DISEASE AND GLIAL CELLS

The brain is composed of many different types of cells, not just neurons.



WHAT IS THE ROLE OF GLIAL CELLS IN PD?

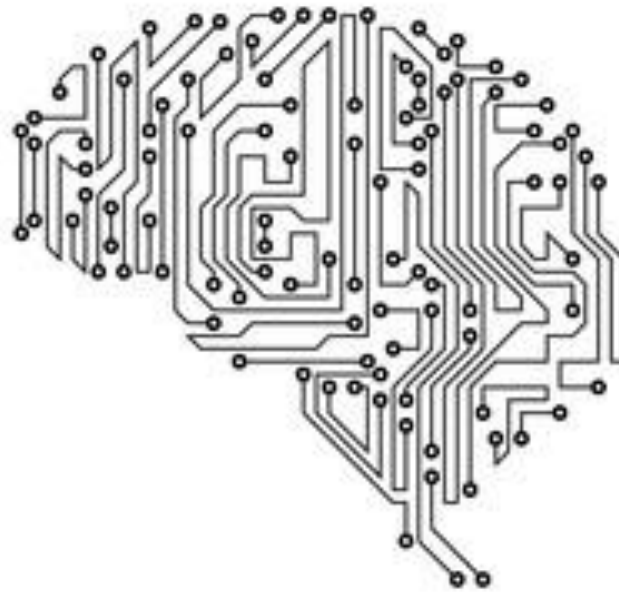


Dr. Abby Olsen, Brigham and Women's Hospital, Boston, MA

Objective:

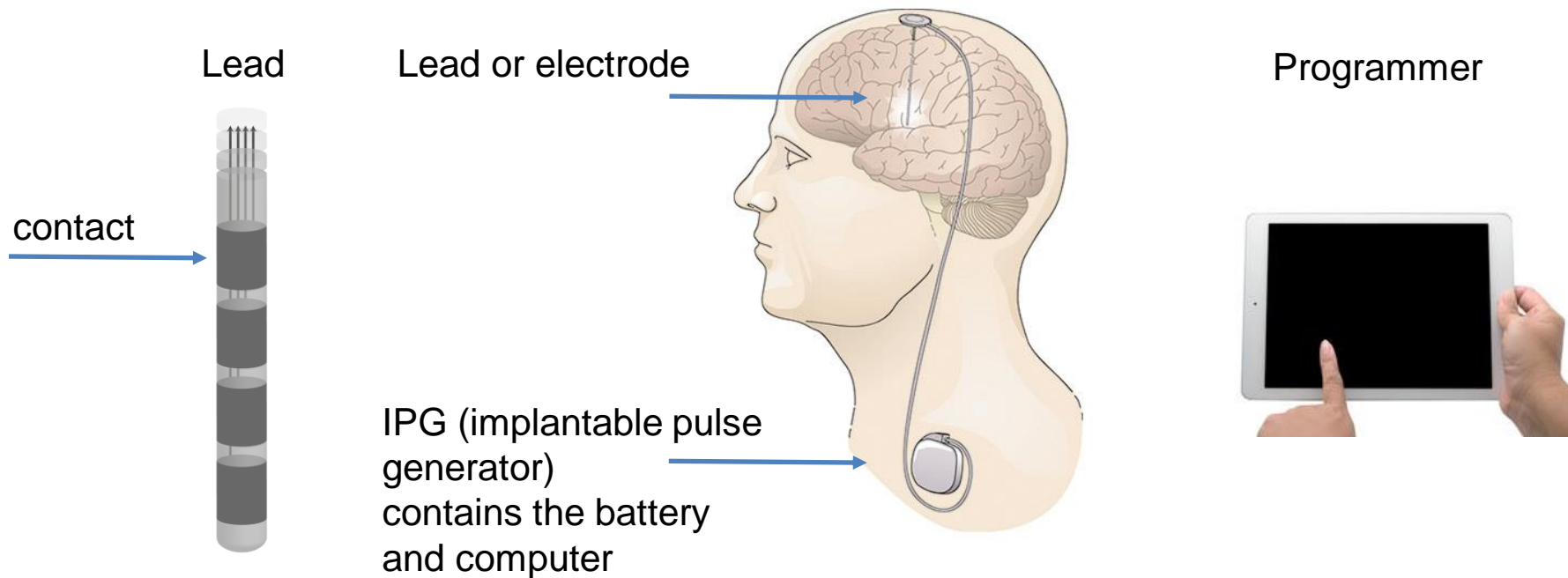
To understand the role of the glial cells in Parkinson's disease in order to eventually develop glial-based therapies for the disease.

PARKINSON'S DISEASE AND BRAIN CIRCUITRY



PARKINSON'S DISEASE AND DEEP BRAIN STIMULATION (DBS)

- DBS is a neurosurgical procedure in which electrodes are placed deep within the brain to deliver electrical impulses to brain structures in order to improve PD symptoms
- The electrodes are connected to an implantable pulse generator (IPG) in the chest, which can be programmed remotely



IS THERE A BETTER WAY TO PERFORM DEEP BRAIN STIMULATION?

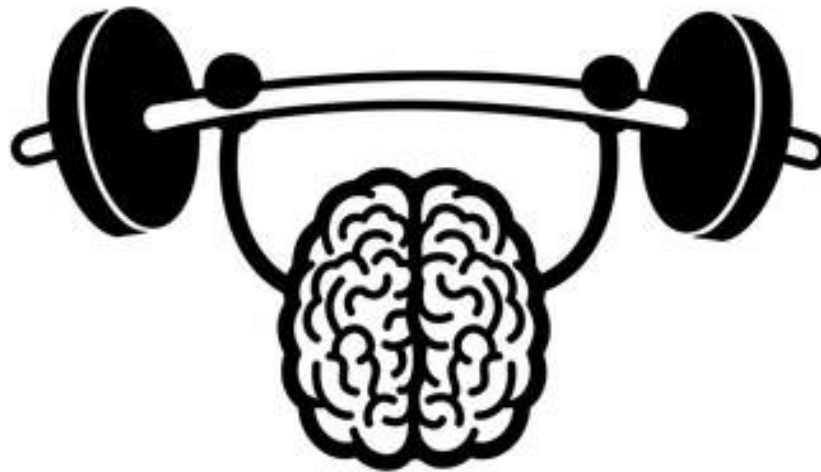


Dr. Enrico Opri, Emory University, Atlanta, GA

Objective:

To investigate the use of Deep Brain Stimulation (DBS) Local Evoked Potentials (DLEPs), brain signals that can be used to guide where the DBS electrode placement during surgery. If these signals are used to guide placement, DBS could be performed more accurately under anesthesia.

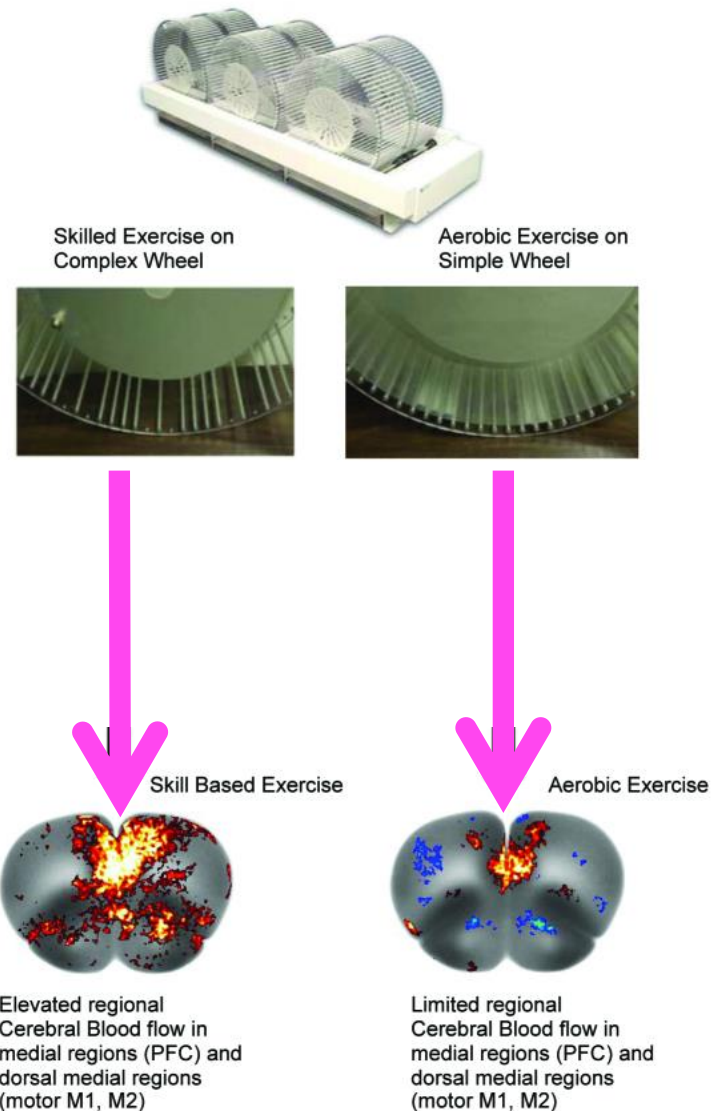
PARKINSON'S DISEASE AND EXERCISE



WHAT DOES EXERCISE DO?

J Hum Kinet. 2016
Sep 1; 52: 35–51.

Learn a new motor skill!



PARKINSON'S DISEASE AND EXERCISE: SPARX STUDY

128 early PD patients, not on dopamine medications, were enrolled in either:

- High-intensity treadmill exercise (4 days per week, 80%-85% maximum heart rate)
- moderate-intensity treadmill exercise (4 days per week, 60%-65% maximum heart rate)
- wait-list control

After 6 months, high intensity group had a 0.3 point change in their United Parkinson's Disease Rating Scale. Control group had a 3 point change ([JAMA Neurol.](#) 2018 Feb; 75(2): 219–226)



[JAMA Neurol.](#) 2018 Feb; 75(2): 219–226.

Published online 2017 Dec 11. doi: [10.1001/jamaneurol.2017.3517](https://doi.org/10.1001/jamaneurol.2017.3517)

PMCID: PMC5838616

PMID: [29228079](https://pubmed.ncbi.nlm.nih.gov/29228079/)

Effect of High-Intensity Treadmill Exercise on Motor Symptoms in Patients With De Novo Parkinson Disease
A Phase 2 Randomized Clinical Trial

WHY IS EXERCISE GOOD FOR THE PD BRAIN?



Dr. Constanza Cortes, University of Alabama, Birmingham, AL

Objective:

To understand the molecular underpinnings of the neuroprotective effects of exercise by determining the effects of exercise on PD pathology in the mouse brain.

PAST RESEARCH ACCOMPLISHMENTS

Novel Variants in *LRRK2* and *GBA* Identified in Latino Parkinson Disease Cohort Enriched for Caribbean Origin

Karen Nuytemans^{1,2*}, Farid Rajabli¹, Parker L. Bussies¹, Katrina Celis¹, William K. Scott^{1,2}, Carlos Simon³, Cornelia C. Luca³, Angel Vinuela⁴, Margaret A. Pericak-Vance^{1,2} and Jeff M. Vance^{1,2}

RESEARCH ARTICLE

Severity-Dependent Effects of Parkinson's Disease on Perceptual Visual and Vestibular Heading

Sinem Balta Beylergil, PhD,^{1,2} Mikkel Petersen, PhD,^{1,2} Palak Gupta, MS,¹ Mohamed Elkasaby, MD,^{4,5} Camilla Kilbane, MD,¹ and Aasef G. Shaikh, MD, PhD^{1,2,*}

¹Department of Biomedical Engineering, Case Western Reserve University, Cleveland, Ohio, USA

²National VA Parkinson Consortium Center, Neurology Service, Daroff-Dell'Osso Ocular Motility and Vestibular Laboratory, Louis Stokes Cleveland VA Medical Center, Cleveland, Ohio, USA

³Department of Clinical Medicine, Center of Functionally Integrative Neuroscience, Aarhus University, Aarhus, Denmark

⁴Department of Neurology, Case Western Reserve University, Cleveland, Ohio, USA

⁵Movement Disorders Center, Neurological Institute, University Hospitals, Cleveland, Ohio, USA

Journal of
Neurochemistry

JNC

The Official Journal of
the International Society
for Neurochemistry



ORIGINAL ARTICLE

Cytisine is neuroprotective in female but not male 6-hydroxydopamine lesioned parkinsonian mice and acts in combination with 17- β -estradiol to inhibit apoptotic endoplasmic reticulum stress in dopaminergic neurons

Sara M. Zarate, Gauri Pandey, Sunandhini Chirukuri, Jose A. Garcia, Brittany Cude, Shannon Storey, Nihal A. Salem, Eric A. Bancroft, Michelle Cook, Rahul Srinivasan✉

First published: 23 December 2020 | <https://doi.org/10.1111/jnc.15282>

ARTICLE OPEN

Sex specific cognitive differences in Parkinson disease

Tyler Harrison Reekes^{1,2}, Christopher Ian Higginson^{1,3}, Christina Raye Ledbetter^{2,4}, Niroshan Sathivadivel^{2,5}, Richard Matthew Zweig⁵ and Elizabeth Ann Disbrow^{1,2,*}✉

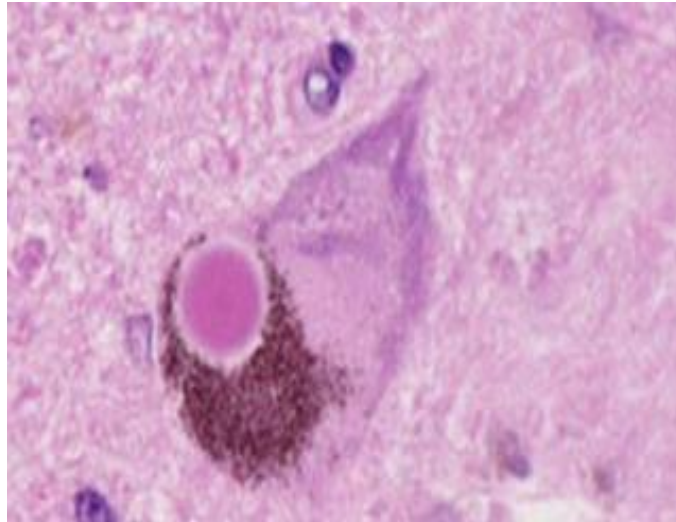
MINI REVIEW
published: 15 January 2020
doi: 10.3389/fpsyg.2019.01494



Pedunculopontine Nucleus Degeneration Contributes to Both Motor and Non-Motor Symptoms of Parkinson's Disease

Nicole Elaine Chambers, Kathryn L. Latta and Christopher Bishop*

PARKINSON'S DISEASE AND ALPHA-SYNUCLEIN



VIVEK UNNI – COTZIAS 2016



Vivek Unni, MD, PhD

- Parkinson Center of Oregon, Department of Neurology and Jungers Center for Neurosciences Research, Oregon Health & Science University, Portland, OR
- Project: *Molecular mechanisms of Lewy body pathology-associated cell death in Parkinson's disease.*
- The study demonstrated that alpha-synuclein plays a role in repair of damaged DNA. This is a brand-new role for alpha-synuclein and can broaden our understanding about what might happen to cause PD

SCIENTIFIC REPORTS

OPEN

Alpha-synuclein is a DNA binding protein that modulates DNA repair with implications for Lewy body disorders

Received: 24 October 2018

Accepted: 12 July 2019

Published online: 29 July 2019

Allison J. Schaser¹, Valerie R. Osterberg¹, Sydney E. Dent¹, Teresa L. Stackhouse¹, Colin M. Wakeham², Sydney W. Boutros³, Leah J. Weston¹, Nichole Owen⁴, Tamily A. Weissman⁵, Esteban Luna⁶, Jacob J. Winkler⁷, Kevin C. Esler⁸, Amanda K. McCullough^{4,7}, Randall L. Woltjer⁹ & Vivek K. Unni^{1,9}

Sci Rep. 2019 Jul 29;9(1):10919. doi: 10.1038/s41598-019-47227-z

PARKINSON'S DISEASE AND GENETICS



KAREN NUYTEMANS – RESEARCH GRANT 2019



Karen Nuytemans, PhD

- John P. Hussman Institute for Human Genomics and John T. Macdonald Department of Human Genetics, University of Miami, Miller School of Medicine, Miami, FL
- Project: *Genetic factors for Parkinson Disease in Hispanics*
- This study identified new mutations in LRRK2 unique to the Latino PD population.



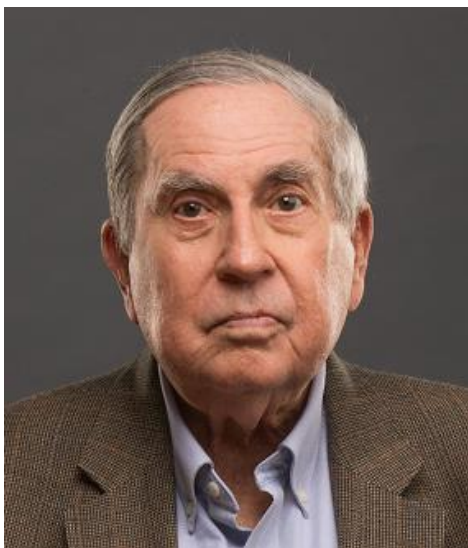
BRIEF RESEARCH REPORT
published: 12 November 2020
doi: 10.3389/fneur.2020.573733

Novel Variants in *LRRK2* and *GBA* Identified in Latino Parkinson Disease Cohort Enriched for Caribbean Origin

Karen Nuytemans^{1,2*}, Farid Rajabli¹, Parker L. Bussies¹, Katrina Celis¹, William K. Scott^{1,2}, Carlos Singer³, Giovanni C. Luca³, Angel Vinuela⁴, Margaret A. Pericak-Vance^{1,2} and Jeff M. Vance^{1,2}

Frontiers in Neurology. 2020. doi: 10.3389/fneur.2020.573733

WHAT ARE THE DIFFERENCES BETWEEN MEN AND WOMEN WHO HAVE PD?



VS



ELIZABETH DISBROW – RESEARCH GRANT 2018



Elizabeth Disbrow, PhD

- Louisiana State University Health Sciences Center, Shreveport, LA
- Project: *Diversity in Biomarker Discovery*
- This study showed that men with PD had significantly greater executive and processing speed impairments compared to women despite no differences in demographic variables or other measures of disease severity.

npj | Parkinson's Disease

www.nature.com/npjparkd

ARTICLE OPEN



Sex specific cognitive differences in Parkinson disease

Tyler Harrison Reekes^{1,2}, Christopher van Hegghebaert³, Christina Raye Ledbetter^{2,4}, Niroshan Sathivadivel^{2,5}, Richard Matthew Zweig¹ and Elizabeth Ann Disbrow^{1,2,5}

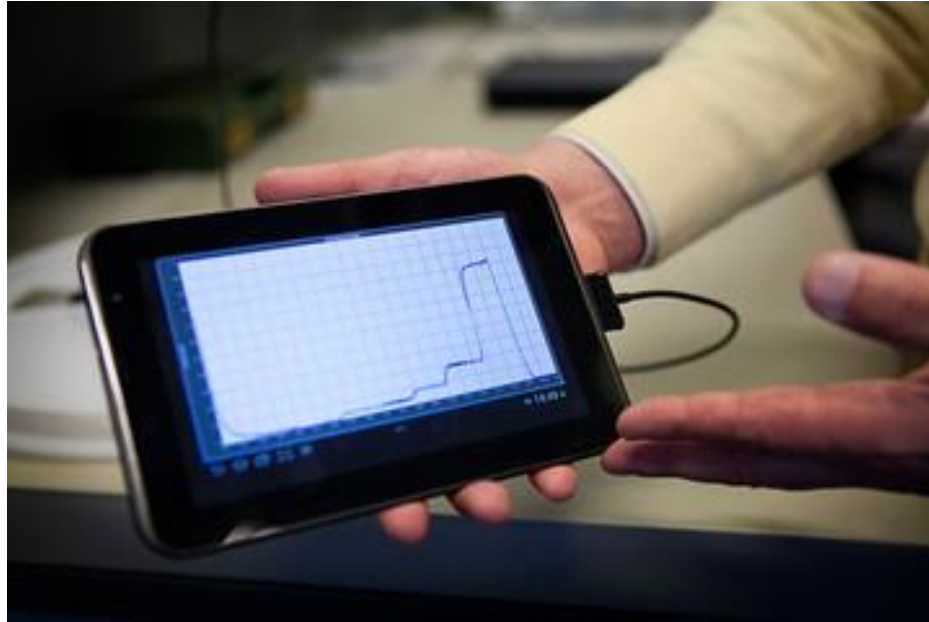
npj Parkinson's Disease. 2020. 6:7 ; <https://doi.org/10.1038/s41531-020-0109-1>



AMERICAN
PARKINSON DISEASE
ASSOCIATION

Strength in optimism. Hope in progress.

PARKINSON'S DISEASE AND DIGITAL BIOMARKERS



YUANFANG GUAN – RESEARCH GRANT 2018



Yuanfang Guan, PhD

- University of Michigan, Ann Arbor, MI
- Project: *Digital Biomarkers for Monitoring Parkinson's Disease (PD)*
- This study developed an algorithm to extract biomarkers of PD from mobile phone accelerometer and gyroscope data

Article

Deep Learning Identifies Digital Biomarkers for Self-Reported Parkinson's Disease

Hanrui Zhang,¹ Kaiwen Deng,¹ Hongyang Li,¹ Roger L. Albin,³ and Yuanfang Guan^{1,4,5}

¹Department of Computational Medicine and Bioinformatics, University of Michigan Medical School, Ann Arbor, MI 48109, USA

²Department of Neurology, University of Michigan Medical School, Ann Arbor, MI, USA

³Neurology Service & GRECC, VAAHS, Ann Arbor, MI 48109, USA

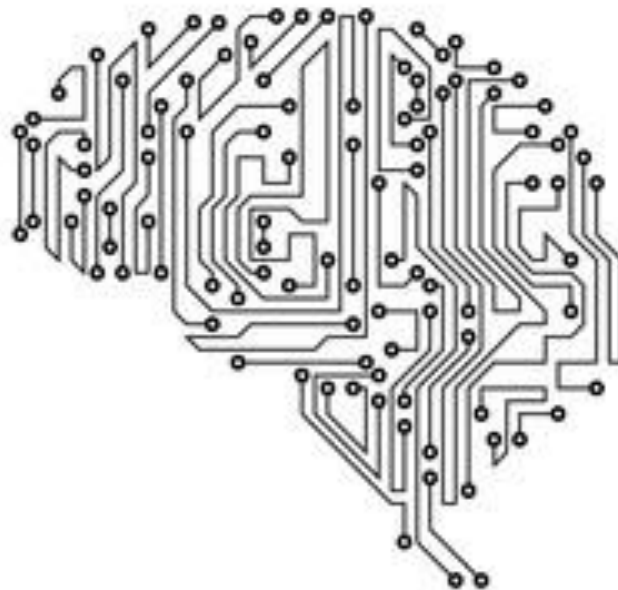
⁴Department of Internal Medicine, University of Michigan Medical School, Ann Arbor, MI 48109, USA

⁵Lead Contact

*Correspondence: gyuanfan@umich.edu

<https://doi.org/10.1016/j.patter.2020.100042>

PARKINSON'S DISEASE AND BRAIN CIRCUITRY



MALORIE HACKER – RESEARCH GRANT 2018



Mallory Hacker, PhD

- Vanderbilt University Medical Center, Nashville, TN
- Project: *Investigating Long-Term Clinical Outcomes of Subthalamic Nucleus Deep Brain Stimulation (DBS) in Early Stage Parkinson's Disease*
- Participants in the early STN DBS + optimal medical therapy group did better at 5 years than those who received optimal medical therapy alone.

ARTICLE CLASS OF EVIDENCE

Deep brain stimulation in early-stage Parkinson disease

Five-year outcomes

Mallory L. Hacker, PhD, MSCI, Max Turchan, MS, Lauren E. Heusinkveld, BS, Amanda D. Currie, MD, Sarah H. Wilson, BS, Anna L. Molinari, JD, Peter E. Konrad, MD, PhD, Thomas L. Davis, MD, Fenna T. Phibbs, MD, MPH, Peter Hedera, MD, PhD, Kevin R. Cannard, MD, Li Wang, MS, and David Charles, MD

Correspondence

Dr. Hacker
Mallory.Hacker@vumc.org

Neurology® 2020;95:e393-e401. doi:10.1212/WNL.0000000000009946

AMERICAN PARKINSON DISEASE ASSOCIATION

TODAY'S GUEST SPEAKERS



Brian Daniels, PhD Vikram Khurana, MD, PhD Karen Nuytemans, PhD Bonnie Wong, PhD

apda

Meet the Researchers: APDA Research Roundtable

971 views • Streamed live on Feb 24, 2021

18 1 SHARE SAVE ...

Virtual Research Roundtable



NEUROINFLAMMATION IN PARKINSON'S DISEASE: AN INTERVIEW WITH ADPA RESEARCHER, DR. EDWARD GRIFFIN



READ MORE >

Dr. Edward Griffin researches the relationship between neuroinflammation and Parkinson's disease. Today we introduce you to Dr. Edward Griffin, a post-doctoral fellow at the University of Alabama at Birmingham. He is the recipient of a 2019-2020 APDA Post-doctoral fellowship and is currently studying the role of inflammation in the development of Parkinson's disease (PD). We asked [...]

APDA-FUNDED BREAKTHROUGHS IN PARKINSON'S RESEARCH



The year 2021 marks the 60th Anniversary of APDA, which was founded in 1961. This milestone has given us the chance to look back at our accomplishments over the past 60 years and reflect on what we have achieved – and also what we have yet to achieve. One of the key pillars of APDA's mission [...]

READ MORE >

WHERE ARE THEY NOW? APDA'S RESEARCH SUCCESS CONTINUED



READ MORE >

Q & A with APDA research grant recipients Since 1961, APDA has been a funding partner in many major scientific breakthroughs and has awarded more than \$51 million in research grants to date. APDA funds individual research grants and fellowships to scientists performing innovative Parkinson's disease (PD) research. Grants are awarded through a competitive application [...]

AMERICAN PARKINSON DISEASE ASSOCIATION

To learn more about what we fund, visit us at
apdaparkinson.org/research

AMERICAN PARKINSON DISEASE ASSOCIATION

APDA is here for you!

To find out more about our programs and services:

Toll-free: 800-223-2732

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