

# Establishing the Frontotemporal Dementia Research Center (FTDRC)

Fen-Biao Gao, PhD

Professor

Governor Paul Cellucci Chair in Neuroscience Research

RNA Therapeutics Institute

UMass Chan Medical School

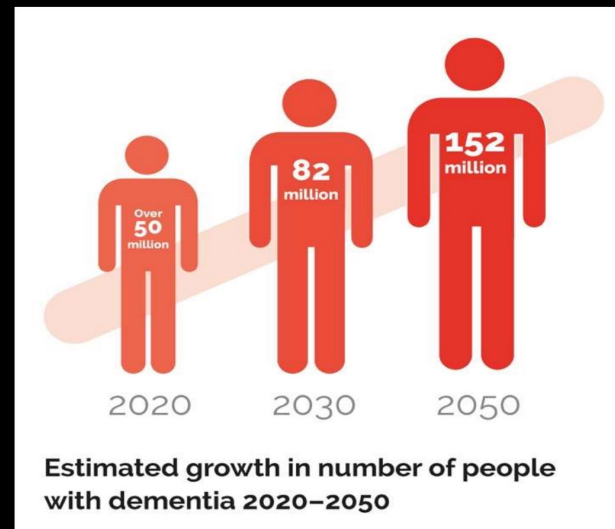
*“There's really realistic hope that we will get a breakthrough and unlock the mystery of ALS and even other neurodegenerative diseases like Alzheimer's. So it's a very promising time.”*

--Governor Paul Cellucci (1948–2013)

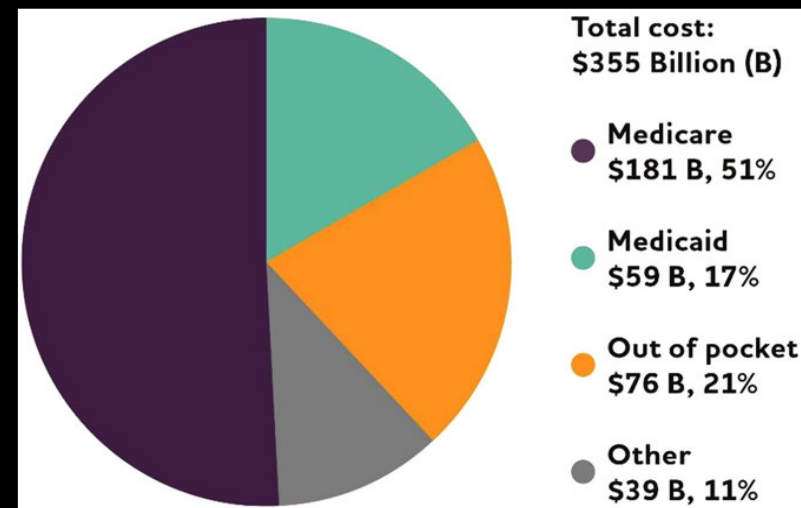


UMass Chan Medical School Chancellor Michael Collins,  
Dr. Robert Brown and the late Gov. Paul Cellucci in 2011

Alzheimer disease (AD) and AD-related dementias (ADRDs) such as frontotemporal dementia (FTD), vascular dementia, dementia with Lewy bodies, and others (such as after a stroke and HIV infection) are a major health challenge in the 21<sup>st</sup> century.

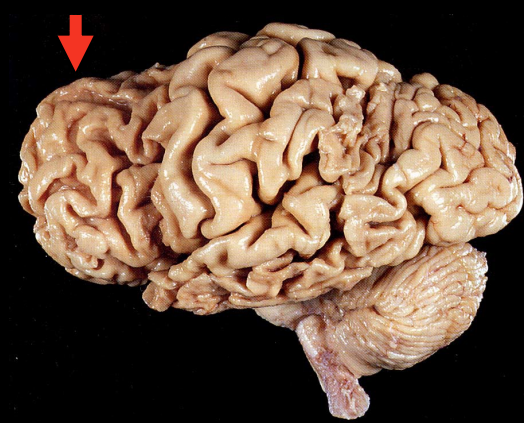


“For FY 2024, NIH estimates the overall funding needed for AD and ADRD totaling \$3.87 billion.”



(Alzheimer’s Association, 2023)

# Frontotemporal Dementia (FTD)



- One of the major Alzheimer's disease-related dementias (ADRDs), and the most common dementia for people under age 60.
- Refers to a group of brain disorders including behavioral variant FTD (bvFTD), primary progressive aphasia (PPA), corticobasal syndrome (CBS), progressive supranuclear palsy (PSP), FTD-motor neuron disease (FTD-MND).
- Progressive deficits in cognitive behaviors (such as social disinhibition and loss of empathy) and/or language, as well as personality changes.
- Affecting 60,000 people in the US, 40% of FTD cases are familial.
- Some genetic, pathological and clinical overlaps with ALS.



# Dementias and Frequency of Criminality

<u>Clinical Diagnosis</u>	<u>Frequency of Criminality</u>
Behavioral Variant FTD	37.4%
Vascular Dementia	14.8%
Alzheimer's Disease	7.7%
Mild Cognitive Impairment	3.3%

Thus, FTD is not only a medical issue but also a social issue.

(Liljegren et al., *JAMA Neurol.* 2015)

## **Vision for the FTD Research Center**

Within 5-10 years, build the Center into a leading place in the nation and the world for FTD (and related neurodegenerative disorders) research and RNA-based therapeutic development.

## My Qualifications

1990–1995	Duke University Medical Center, USA	PhD	RNA biology
1995–1997	University of College London, UK	Postdoc	Neurobiology
1997–2000	University of California, San Francisco	Postdoc	Neurobiology
07.2000–05.2006	Assistant Investigator, Gladstone Institute of Neurological Disease (GIND) Assistant Professor, Department of Neurology, UCSF		
06.2006–02.2010	Associate Investigator, GIND; Associate Professor of Neurology, UCSF		
02.2010–present	Professor, Department of Neurology, UMass Chan Medical School		
08.2023–present	Professor, RNA Therapeutics Institute, UMass Chan Medical School		

# My Qualifications

>130 papers. Representative papers in FTD/ALS research since I came to UMass Chan in 2010:

## As the (co) corresponding author

Almeida et al., *Cell Reports* 2012  
Lu et al., *Mol. Cell* 2013  
Gascon et al., *Nat. Med.* 2014  
Yang et al., *Acta Neuropathol.* 2015  
Tran et al., *Neuron* 2015  
Freibaum et al., *Nature* 2015  
Lopez-Gonzalez et al., *Neuron* 2016  
Gao et al., *EMBO J.* 2017  
Gao et al., *Cell* 2017  
Yuva-Aydemir et al., *Trends Neurosci.* 2018  
Lopez-Gonzalez et al., *PNAS* 2019  
Yuva-Aydemir et al., *Nat. Commun.* 2019  
Choi et al., *Nat. Neurosci.* 2019  
Krishnan et al., *Acta Neuropathol.* 2020  
Lu et al., *Acta Neuropathol.* 2021  
van't Spijker et al., *RNA* 2021  
Loveland et al., *Nat. Commun.* 2022  
Krishnan et al., *Nat. Commun.* 2022  
Jun et al., *Autophagy* 2023  
Lee et al., *Neuron* 2023

## As a collaborator

Boxer et al., *Alzheimer's and Dementia* 2013a  
Boxer et al., *Alzheimer's and Dementia* 2013b  
Filiano et al., *J. Neurosci.* 2013  
West et al., *J. Cell Biol.* 2015  
Peters et al., *Neuron* 2015  
Burguete et al., *Elife* 2015  
Woehlbier et al., *EMBO J.* 2016  
Kramer et al., *Science* 2016  
Yin et al., *Cell Reports* 2017  
Markmiller et al., *Cell* 2017  
Chen et al., *Neuron* 2019  
Li et al., *Cell Reports* 2020  
Maor-Nof et al., *Cell* 2021  
Sonobe et al., *Nat. Commun.* 2021  
Arredondo et al., *Neuron* 2022  
Meijboom et al., *Nat. Commun.* 2022  
Hung et al., *Cell* 2023  
Philips et al., *Neuron* 2023  
Fang et al., *Front. Neurosci.* In press.

(With UMass Chan collaborators: Zhiping, Andrei, Jon, Bob, Joel, Neal, Miguel, Michael Green.....)



## My Qualifications

Leadership positions and professional services:

- Vice Chair of Research, Department of Neurology
- Chair of Tenure Committee, Department of Neurology
- Member of the University Tenure Committee
  
- Co-organizer of “RNA Metabolism in Neurological Disease” conferences 5 times
- Co-organizer of other meetings
- Member of the NIH Cellular Molecular Neurodegeneration study section (2019–2023)
- Member of the NIH F03A Fellowship Study Section (2011–2015)
- Member of the NIH Synapse and Trafficking study section (2007–2011)
- Senior Editor, Brain Research (2011–2021)

2023–present: Member of the Medical Advisory Council, Association for FTD

2023–present: Member of the Postdoctoral Fellowship Review Panel, King Trust

2020–present: Member of the SAB, NINDS Human Cell and Data Repository

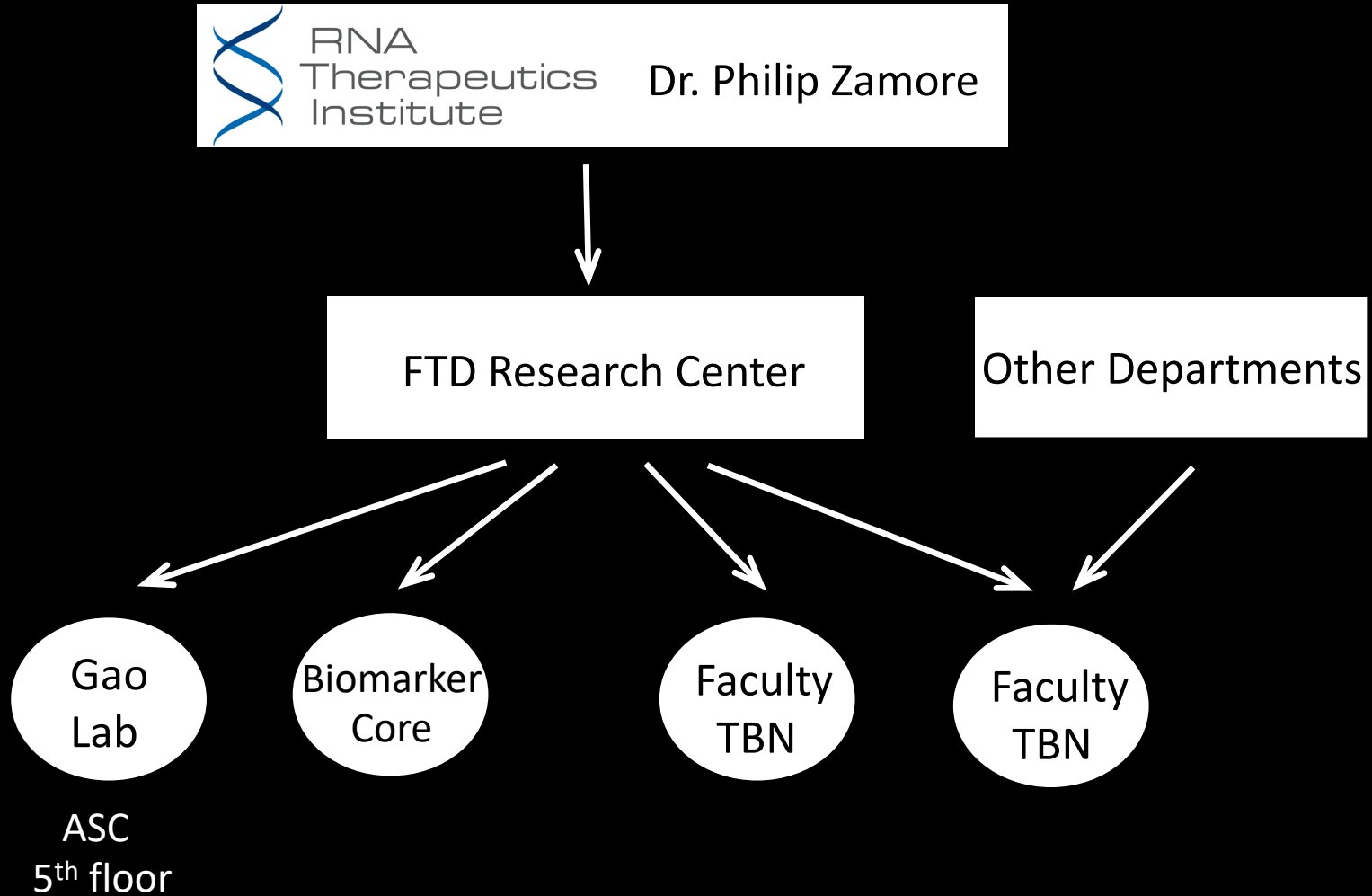
2018–present: Section Editor, *Translational Neurodegeneration* (IF>12)

2017–present: Member of the Research Advisory Council, Muscular Dystrophy Association

2015–present: Member of the Grant Review Panel, ALS Association

2015–present: Member of the Grant Review Panel, Association for FTD

# Organizations



## Recruitments

2-3 highly talented, driven, and diverse junior faculty in 4-6 years.

- Dysregulation of RNA metabolism in FTD.
- Synaptic and circuit mechanisms underlying abnormal behaviors in FTD.
- Neuropathologist (MD/PhD), dementia brain bank, scRNA-seq, AI etc.
- Other cutting-edge research areas.
- Focus on people (next Craig Mello, Philip Zamore.....), not projects.

## Fundings

- Private donations.
- Foundation grants (Alzheimer's Association, Association for FTD, etc.).
- NIA FTD program grant (P01) (Due January 25, 2024, \$1.5–1.7 million direct/year).  
*“Frontotemporal Dementia: From Cryo-EM, Neural Mechanisms to RNA Therapy”*
- Campus-wide NIH/NINDS T32 training grant on FTD (Due May 25, 2024, or 2025).
- Future program grants and multi-PI R01 grants, once the Center grows.

## Services to RTI, UMass Chan and Beyond

- Contribute to the exciting, respectful and inclusive research and teaching environment at the RTI.
- Use my expertise in grant writing to help junior faculty and trainees in the Center and the Department to obtain external fundings.
- Foster scientific excellent in the FTD Research Center and encourage collaborations with other departments.
- Bridge the interactions between basic scientists and clinicians in dementia field across the UMass Chan campus.
- Engage with the public in Massachusetts and promote dementia awareness. (September 24–October 1, World FTD Awareness Week)

## **Vision for the FTD Research Center**

Within 5-10 years, build the Center into a leading place in the nation and the world for FTD (and related neurodegenerative disorders) research and RNA-based therapeutic development.