# Charles D. Chen

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#### Education

#### Washington University in St. Louis

2018-2023

Ph.D. in Imaging Science M.S. in Electrical Engineering

## Washington University in St. Louis

2011-2015

B.A. in Biochemistry

# Experience -

### Washington University in St. Louis

2017-2023

Research assistant (PI: Tammie L.S. Benzinger)

- Co-founded the PAINT committee and established its imaging-pathology pipeline for the Knight ADRC and DIAN
- Established a computational approach for finding optimal ROIs and cutoffs for AD biomarkers in the Knight ADRC and DIAN
- Established a computational approach for estimating drug effects from imaging and pathology in the DIAN-TU-001 clinical trial

University of Oxford 2015-2015

Visiting scholar (PI: Stamatios N. Sotiropoulos)

• Established an imaging-pathology approach for developing new HCP tractography algorithms

#### Washington University in St. Louis

2012-2015

Research assistant (PI: David C. Van Essen)

• Established an imaging-pathology approach for estimating biases in HCP tractography algorithms

#### Skills —

#### Computational methods

Computer vision, Data science, High-performance computing, Machine learning, Optimization, Statistics

#### **Programming languages**

Bash, MATLAB, Python, R

#### Team management

Onboarded and mentored trainees in data acquisition, data analysis, programming, statistics, and manuscript writing

## Honors and awards -

2023	Poletsky Award	\$2,000
2021	McKelvey School of Engineering Dean's Scholarly Award	\$5,000
2020	Knight ADRC T32 Fellowship	\$25,836
2020	HAI Travel Scholarship Winner and Young Investigator Award Finalist	\$1,000
2018	NSF Graduate Research Fellowship Program	\$138,000
2013	HHMI Summer Undergraduate Research Fellowship	\$4,000
2011	National Merit Scholarship	\$2,500

## **Service**

2022-2023	Quantitative Imaging Biomarkers Alliance (QIBA) Tau Profile Committee	Assistant to the co-chairs
2022-2023	Diversity, Equity, and Inclusion (DEI) and Well-being Steering Committee	
2022-2023	Imaging Science Student Council Executive Board	Co-founder
2020-2021	Nature Reviews Neurology	Ad hoc reviewer
2017-2023	Pathology And Imaging Neurodegeneration and Tau (PAINT) Committee	Co-founder

# **Publications**

- Accepted Kanta Horie, Gemma Salvadó, Nico Barthélemy, Shorena Janelidze, Yan Li, Yingxin He, Benjamin Saef, **Charles D Chen**, Hong Jiang, Olof Standberg, Alexa Pichet Binette, Sebastian Palmqvist, Chihiro Sato, Brian A Gordon, Tammie LS Benzinger, David M Holtzman, John C Morris, Niklas Mattsson-Carlgren, Erik Stomrud, Rik Ossenkoppele, Suzanne E Schindler, Oskar Hansson, and Randall J Bateman. CSF MTBR-tau243 is a specific biomarker of tau tangle pathology in Alzheimer's disease. *Nature Medicine*
- Stephanie A Schultz, Zahra Shirzadi, Aaron P Schultz, Lei Liu, Colleen D Fitzpatrick, Eric McDade, Alan Renton, Bianca Esposito, Nelly Joseph-Mathurin, Carlos Cruchaga, **Charles D Chen**, Alison Goate, Ricardo Francisco Allegri, Tammie LS Benzinger, Sarah Berman, Helena C Chui, Anne M Fagan, Martin R Farlow, Nick C Fox, Brian A Gordon, Neill R Graff-Radford, Jason J Hassenstab, Bernard J Hanseeuw, Anna Hofmann, Clifford R Jack Jr, Mathias Jucker, Celeste M Karch, Robert A Koeppe, Jae-Hong Lee, Allan I Levey, Johannes Levin, Ralph N Martins, Hiroshi Mori, John C Morris, James Noble, Richard J Perrin, Pedro Rosa-Neto, Stephen P Salloway, Raquel Sanchez-Valle, Peter R Schofield, Chengjie Xiong, Keith A Johnson, Randall J Bateman, Reisa A Sperling, Jasmeer P Chhatwal, and the Dominantly Inherited Alzheimer Network Investigators. Location of pathogenic variants in PSEN1 impacts progression of cognitive, clinical, and neurodegenerative measures in autosomal-dominant AD. *Aging Cell*. http://doi.org/10.1111/acel.13871
- Charles Chen\*, Maria Rosana Ponisio\*, Jordan Lang, Shaney Flores, Suzanne Schindler, Anne Fagan, John Morris, and Tammie Benzinger. Comparing tau PET visual interpretation with tau PET quantification, CSF biomarkers, and longitudinal clinical assessment. *Journal of Alzheimer's Disease*. https://doi.org/10.3233/JAD-230032
- Charles D Chen\*, Austin McCullough\*, Brian A Gordon\*, Nelly Joseph-Mathurin, Shaney Flores, Nicole S McKay, Diana Hobbs, Russ Hornbeck, Anne M Fagan, Carlos Cruchaga, Alison M Goate, Richard J Perrin, Guoqiao Wang, Yan Li, Xinyu Shi, Chengjie Xiong, Michael J Pontecorvo, Gregory Klein, Yi Su, William E Klunk, Clifford Jack, Robert Koeppe, B Joy Snider, Sarah B Berman, Erik D Roberson, Jared Brosch, Ghulam Surti, Ivonne Z Jiménez-Velázquez, Douglas Galasko, Lawrence S Honig, William S Brooks, Roger Clarnette, David Wallon, Bruno Dubois, Jérémie Pariente, Florence Pasquier, Raquel Sanchez-Valle, Sergey Shcherbinin, Ixavier Higgins, Ilke Tunali, Colin L Masters, Christopher H van Dyck, Mario Masellis, Robin Hsiung, Serge Gauthier, Steve Salloway, David B Clifford, Susan Mills, Charlene Supnet-Bell, Eric McDade, Randall J Bateman, Tammie LS Benzinger, and for the DIAN-TU Study Team. Longitudinal head-to-head comparison of <sup>11</sup>C-PiB and <sup>18</sup>F-florbetapir PET in a Phase 2/3 clinical trial of anti-amyloid-β monoclonal antibodies in dominantly inherited Alzheimer disease. *European Journal of Nuclear Medicine and Molecular Imaging*. https://doi.org/10.1007/s00259-023-06209-0
- Nicole S McKay, Brian A Gordon, Russ C Hornbeck, Aylin Dincer, Shaney Flores, Sarah Keefe, Nelly Joseph-Mathurin, Clifford R Jack, Robert Koeppe, Peter R Millar, Beau M Ances, **Charles D Chen**, Alisha Daniels, Diana A Hobbs, Kelley Jackson, Deborah Koudelis, Parinaz Massoumzadeh, Austin McCullough, Michael L Nickels, Farzaneh Rahmani, Laura Swisher, Qing Wang, Ricardo F Allegri, Sarah B Berman, Adam M Brickman, William S Brooks, David M Cash, Jasmeer P Chhatwal, Gregg Day, Martin R Farlow, Christian IaFougère, Nick C Fox, Michael Fulham, Bernardino Ghetti, Neill Graff-Radford, Takeshi Ikeuchi, William Klunk, Jae-Hong Lee, Ralph Martins, Colin L Masters, Jonathan McConathy, Hiroshi Mori, James M Noble, Gerald Reischl, Christopher Rowe, Stephen Salloway, Raquel Sanchez-Valle, Peter R Schofield, Hiroyuki Shimada, Mikio Shoji, Yi Su, Kazushi Suzuki, Jonathan Vöglein, Igor Yakushev, Carlos Cruchaga, Jason Hassenstab, Celeste Karch, Eric McDade, Richard J Perrin, Chengjie Xiong, John C Morris, Randall J Bateman, Tammie LS Benzinger, and for the Dominantly Inherited Alzheimer Network. Positron emission tomography and magnetic resonance imaging methods and datasets within the Dominantly Inherited Alzheimer Network (DIAN). *Nature Neuroscience*
- Farzaneh Rahmani, Saurabh Jindal, Cyrus A Raji, Wei Wang, Arash Nazeri, Gloria Guzman Perez-Carillo, Michelle M Miller-Thomas, Philipp Garner, Benedicte Marechal, Amit Shah, Mathis Zimmermann, **Charles D Chen**, Sarah Keefe, Pamela LaMontagne, and Tammie LS Benzinger. Validity assessment of an automated brain morphometry tool using patients with a de novo memory complaint. *American Journal of Neuroradiology*. https://doi.org/10.3174/ajnr.A7790

- Anna H Boerwinkle, Brian A Gordon, Julie Wisch, Shaney Flores, Rachel L Henson, Omar H Butt, Nicole McKay, Charles D Chen, Tammie LS Benzinger, Anne M Fagan, Benjamin L Handen, Bradley T Christian, Elizabeth Head, Mark Mapstone, Michael S Rafii, Sid O'Bryant, Florence Lai, H Diana Rosas, Joseph H Lee, Wayne Silverman, Adam M Brickman, Jasmeer P Chhatwal, Carlos Cruchaga, Richard J Perrin, Chengjie Xiong, Jason Hassenstab, Eric McDade, Randall J Bateman, Beau M Ances, on behalf of the Alzheimer's Biomarker Consortium-Down Syndrome, and the Dominantly Inherited Alzheimer Network. Amyloid PET in genetic causes of Alzheimer disease: Down syndrome versus autosomal dominant. *The Lancet Neurology*. https://doi.org/10.1016/S1474-4422(22)00408-2
- Aylin Dincer, **Charles D Chen**, Nicole S McKay, Lauren N Koenig, Austin McCullough, Shaney Flores, Sarah J Keefe, Stephanie A Schultz, Rebecca L Feldman, Nelly Joseph-Mathurin, Russ C Hornbeck, Carlos Cruchaga, Suzanne E Schindler, David M Holtzman, John C Morris, Anne M Fagan, Tammie LS Benzinger, and Brian A Gordon. APOE ε4 genotype, amyloid-β, and sex interact to predict tau in regions of high APOE mRNA expression. *Science Translational Medicine*. http://doi.org/10.1126/scitranslmed.abl7646
- Guoqiao Wang\*, Yan Li\*, Chengjie Xiong, Eric McDade, David B Clifford, Susan L Mills, Anna M Santacruz, Andrew J Aschenbrenner, Jason Hassenstab, Tammie LS Benzinger, Brian A Gordon, Anne M Fagan, Kelley A Coalier, Jorge J Libre-Guerra, Austin McCullough, Nelly Joseph-Mathurin, **Charles Chen**, Catherine Mummery, Barbara A Wendelberger, Serge Gauthier, Mario Masellis, Karen C Holdridge, Roy Yaari, Saptarshi Chatterjee, John Sims, Paul Delmar, Geoffrey A Kerchner, Tobias Bittner, Carsten Hofmann, Randall J Bateman, and for the DIAN-TU Study Team. Evaluation of dose-dependent treatment effects after mid-trial dose escalation in biomarker, clinical, and cognitive outcomes for gantenerumab or solanezumab in dominantly inherited Alzheimer's disease. Alzheimer's & Dementia: Diagnosis, Assessment & Disease Monitoring. http://doi.org/10.1002/dad2.12367
- Nelly Joseph-Mathurin\*, Jorge J Llibre-Guerra\*, Yan Li, Austin A McCullough, Carsten Hofmann, Jakub Wojtowicz, Ethan Park, Guoqiao Wang, Gregory M Preboske, Qing Wang, Brian A Gordon, **Charles D Chen**, Shaney Flores, Neelum Aggarwal, Sarah B Berman, Thomas D Bird, Sandra E Black, Bret Borowski, William S Brooks, Jasmeer P Chhatwal, Roger Clarnette, Carlos Cruchaga, Anne M Fagan, Martin Farlow, Nick C Fox, Serge Gauthier, Jason Hassenstab, Diana A Hobbs, Karen C Holdridge, Lawrence S Honig, Russ C Hornbeck, Ging-Yuek R Hsiung, Clifford R Jack Jr, Ivonne Z Jimenez-Velazquez, Mathias Jucker, Gregory Klein, Johannes Levin, Michele Mancini, Mario Masellis, Nicole S McKay, Catherine J Mummery, John M Ringman, Hiroyuki Shimada, Joy Snider, Kazushi Suzuki, David Wallon, Chengjie Xiong, Roy Yaari, Eric McDade, Richard J Perrin, Randall J Bateman, Stephen P Salloway, Tammie LS Benzinger, David B Clifford, and for the Dominantly Inherited Alzheimer Network Trials Unit. Amyloid-related imaging abnormalities in the DIAN-TU-001 trial of gantenerumab and solanezumab: lessons from a trial in dominantly inherited Alzheimer disease. *Annals of Neurology*. https://doi.org/10.1002/ana.26511
- Shaney Flores, **Charles D Chen**, Yi Su, Aylin Dincer, Sarah J Keefe, Nicole S McKay, Angela M Paulick, Gloria Guzman Perez-Carillo, Russ C Hornbeck, Manu S Goyal, Andrei G Vlassenko, Sally W Schwarz, Michael L Nickels, Dean F Wong, Zhude Tu, Jonathan McConathy, John C Morris, Tammie Benzinger, and Brian A Gordon. Investigating tau and amyloid tracer skull binding in studies of Alzheimer disease. *Journal of Nuclear Medicine*. https://doi.org/10.2967/jnumed.122.263948
- Patrick H Luckett, **Charlie Chen**, Brian A Gordon, Julie Wisch, Sarah B Berman, Jasmeer P Chhatwal, Carlos Cruchaga, Anne M Fagan, Martin R Farlow, Nick C Fox, Mathias Jucker, Johannes Levin, Colin L Masters, Hiroshi Mori, James M Noble, Stephen Salloway, Peter R Schofield, Adam M Brickman, William S Brooks, David M Cash, Michael J Fulham, Bernardino Ghetti, Clifford R Jack, Jonathan Vöglein, William Klunk, Robert Koeppe, Yi Su, Michael Weiner, Qing Wang, Dan Marcus, Deborah Koudelis, Nelly Joseph-Mathurin, Lisa Cash, Russ Hornbeck, Chengjie Xiong, Richard J Perrin, Celeste M Karch, Jason Hassenstab, Eric McDade, John C Morris, Tammie LS Benzinger, Randall J Bateman, Beau M Ances, and for the Dominantly Inherited Alzheimer Network (DIAN). Biomarker clustering in autosomal dominant Alzheimer disease. *Alzheimer's & Dementia*. https://doi.org/10.1002/alz.052149
- Farzaneh Rahmani, Marina Nguyen, **Charles D Chen**, Nicole McKay, Aylin Dincer, Nelly Joseph-Mathurin, Gengsheng Chen, Jingxia Liu, Hilary LP Orlowski, John C Morris, and Tammie Benzinger. Intracranial internal carotid artery calcification is not predictive of future cognitive decline. *Alzheimer's Research & Therapy*. https://doi.org/10.1186/s13195-022-00972-2

- Charles D Chen, Nelly Joseph-Mathurin\*, Namita Sinha\*, Aihong Zhou\*, Yan Li, Karl Friedrichsen, Austin McCullough, Erin E Franklin, Russ Hornbeck, Brian Gordon, Vijay Sharma, Carlos Cruchaga, Alison Goate, Celeste Karch, Eric McDade, Chengjie Xiong, Randall J Bateman, Bernardino Ghetti, John M Ringman, Jasmeer Chhatwal, Colin L Masters, Catriona McLean, Tammaryn Lashley, Yi Su, Robert Koeppe, Clifford Jack, William Klunk, John C Morris, Richard J Perrin, Nigel J Cairns\*\*, and Tammie LS Benzinger\*\*. Comparing amyloid-β plaque burden with antemortem PiB PET in autosomal dominant and late-onset Alzheimer disease. *Acta Neuropathologica*. https://doi.org/10.1007/s00401-021-02342-y
- Anna H Boerwinkle\*, Julie K Wisch\*, **Charles D Chen**, Brian A Gordon, Omar H Butt, Suzanne E Schindler, Courtney Sutphen, Shaney Flores, Aylin Dincer, Tammie LS Benzinger, Anne M Fagan, John C Morris, and Beau M Ances. Temporal correlation of CSF and neuroimaging in the Amyloid-Tau-Neurodegeneration model of Alzheimer disease. *Neurology*. https://doi.org/10.1212/WNL.000000000012123
- Charles D Chen, Timothy R Holden, Brian A Gordon, Erin E Franklin, Yan Li, Dean W Coble, Hongbo Luo, Randall J Bateman, for the Dominantly Inherited Alzheimer Network (DIAN), for the Dominantly Inherited Alzheimer Network Trials Unit (DIAN-TU), Beau M Ances, Richard J Perrin, Tammie LS Benzinger, Nigel J Cairns, and John C Morris. Ante- and postmortem tau in autosomal dominant and late-onset Alzheimer disease. Annals of Clinical and Translational Neurology. https://doi.org/10.1002/acn3.51237
- Brian A Gordon, Tyler M Blazey, Jon Christensen, Aylin Dincer, Shaney Flores, Sarah Keefe, **Charles Chen**, Yi Su, Eric M McDade, Guoqiao Wang, Yan Li, Jason Hassenstab, Andrew Aschenbrenner, Russ Hornbeck, Clifford R Jack, Jr, Beau M Ances, Sarah B Berman, Jared R Brosch, Douglas Galasko, Serge Gauthier, James J Lah, Mario Masellis, Christopher H van Dyck, Mark Mintun, Gregory Klein, Smiljana Ristic, Nigel J Cairns, Daniel S Marcus, Chengjie Xiong, David M Holtzman, Marcus E Raichle, John C Morris, Randall J Bateman, and Tammie LS Benzinger. Tau PET in autosomal dominant Alzheimer's disease: relationship with cognition, dementia and other biomarkers. *Brain*. https://doi.org/10.1093/brain/awz019
- Qing Wang, Yong Wang, Jingxia Liu, Courtney L Sutphen, Carlos Cruchaga, Tyler Blazey, Brian A Gordon, Yi Su, Charlie Chen, Joshua S Shimony, Beau M Ances, Nigel J Cairns, Anne M Fagan, John C Morris, and Tammie LS Benzinger. Quantification of white matter cellularity and damage in preclinical and early symptomatic Alzheimer's disease. *NeuroImage: Clinical*. https://doi.org/10.1016/j.nicl.2019.101767
- Kaining Zhang, **Charles D Chen**, and Ilya E Monosov. Novelty, salience, and surprise timing are signaled by neurons in the basal forebrain. *Current Biology*. https://doi.org/10.1016/j.cub.2018.11.012
- Michiel Cottaar, Matteo Bastiani, **Charles Chen**, Krikor Dikranian, David Van Essen, Timothy E Behrens, Stamatios N Sotiropoulos, and Saad Jbabdi. A gyral coordinate system predictive of fibre orientations. *NeuroImage*. https://doi.org/10.1016/j.neuroimage.2018.04.040
- Noah M Ledbetter, **Charles D Chen**, and Ilya E Monosov. Multiple mechanisms for processing reward uncertainty in the primate basal forebrain. *Journal of Neuroscience*. https://doi.org/10.1523/jneurosci.1123-16.2016

# Book chapters

David C Van Essen, Saad Jbabdi, Stamatios N Sotiropoulos, **Charles Chen**, Krikor Dikranian, Tim Coalson, John Harwell, Timothy EJ Behrens, and Matthew F Glasser. Mapping connections in humans and non-human primates: aspirations and challenges for diffusion imaging. *Diffusion MRI: From Quantitative Measurement to In vivo Neuroanatomy*. https://doi.org/10.1016/B978-0-12-396460-1.00016-0

## **Conference talks**

- 2021 **Charles D Chen**, Jordan A Lang, Shaney Flores, Suzanne E Schindler, Anne M Fagan, John C Morris, Tammie LS Benzinger, and Maria Rosana Ponisio. Concordance of PET tau visual reads with PET tau quantification and CSF pTau. *Radiological Society of North America*
- Charles D Chen, Timothy R Holden, Brian A Gordon, Erin E Franklin, Richard J Perrin, John C Morris, Randall Bateman, Tammie LS Benzinger, Nigel J Cairns, and for the Dominantly Inherited Alzheimer Network (DIAN). Tauopathy in autosomal dominant and late-onset Alzheimer disease. *Alzheimer's Association International Conference*
- Stamatios N Sotiropoulos, **Charlie Chen**, Krikor Dikranian, Saad Jbabdi, Timothy EJ Behrens, David C Van Essen, and Matthew F Glasser. Comparison of diffusion MRI predictions and histology in the macaque brain. *International Society for Magnetic Resonance in Medicine*

# **Invited talks**

- **Charles D Chen**. Our experience with FDA-approved guidelines for visual interpretation of <sup>18</sup>F-flortaucipir PET for Alzheimer disease. *Knight ADRC Tuesday Seminar*
- **Charles D Chen**. Evaluating PET imaging with visual reads, neuropathology, and longitudinal head-to-head comparisons. *Martinos Center SuperGroup Talk*
- **Charles D Chen**. Longitudinal PET amyloid-β shows radiotracer-specific drug effects in the DIAN-TU-001 trial. Knight ADRC Tuesday Seminar
- **Charles D Chen**. Ante- and postmortem tau in autosomal dominant and late-onset Alzheimer disease. *DIAN-Obs Chalk Talk*