



Data Sharing In DIAN

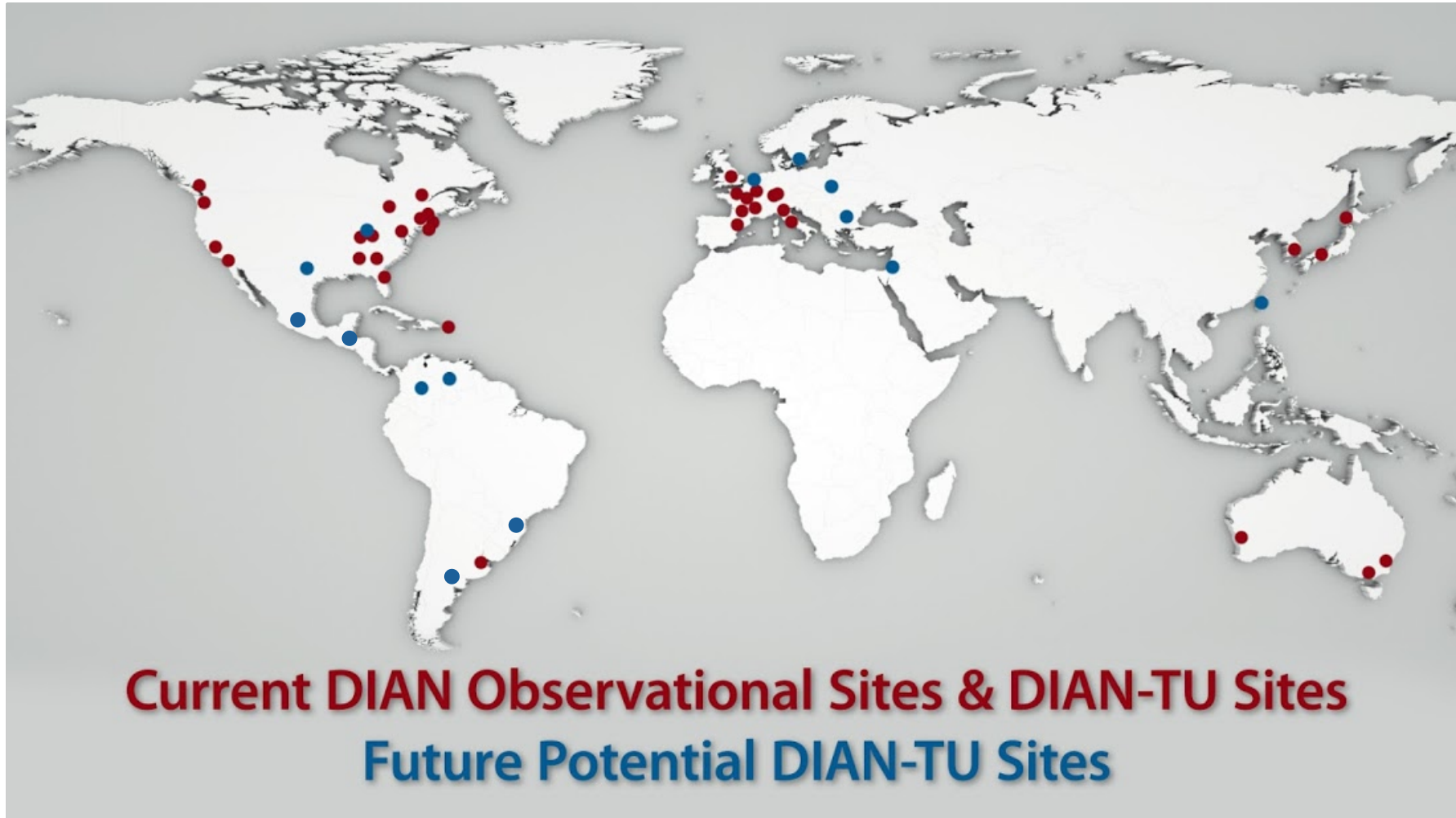
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Associate Director, DIAN Trials Unit
Washington University School of Medicine



DIAN Observational *and* DIAN-TU Trial Sites

The DIAN Obs has currently enrolled over 550 participants.



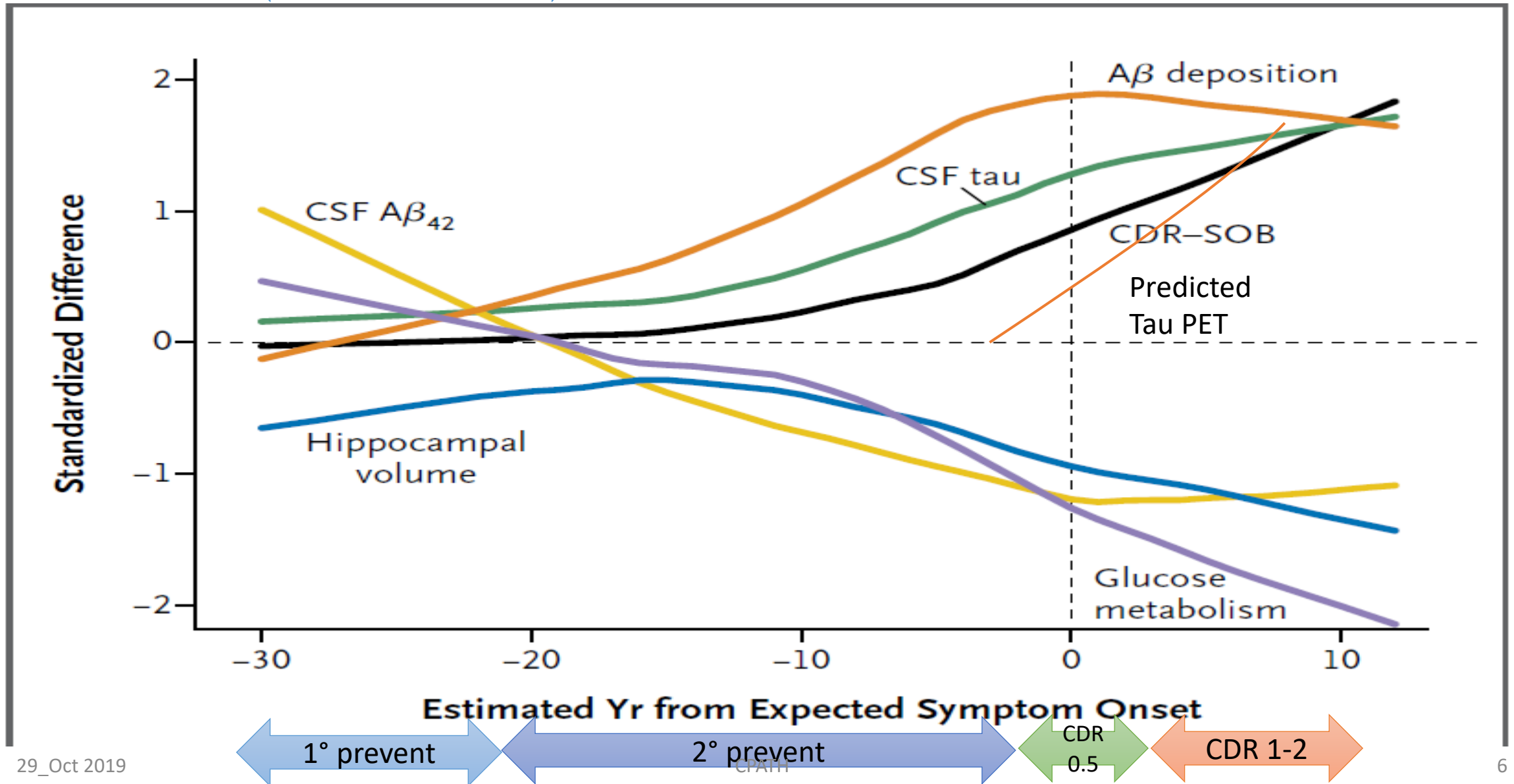
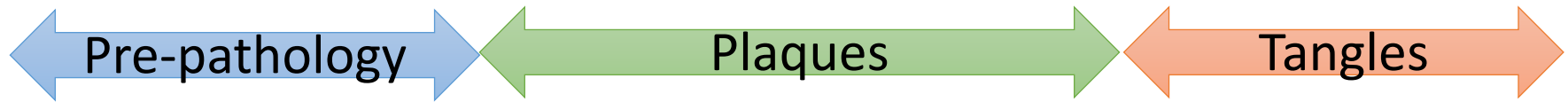
Participant Entry Characteristics at Enrollment

N = 571* (Target 80% Asymptomatic, 20% Symptomatic) (*Table based on 534 participants. 37 Mutations in Process)	Asymptomatic 421 (73.7%) 392 with confirmed mutation status		Symptomatic 146 (25.6%) 140 with confirmed mutation status		
	200 (NC) (51.0%)	192 (MC) (46.2%)	10 (NC) (7.1%)	130 (MC) (92.9%)	
Age, Mean (SD)	41.6 (12.1)	37.2 (9.6)	44.1 (12.8)	47.4 (9.2)	
Gender (% Female)	117 (58.5%)	107 (55.7%)	6 (60.0%)	73 (56.2%)	
Parental Age of Onset, Mean (SD)	47.2 (6.5)	48.5 (7.1)	47.9 (6.2)	45.6 (8.7)	
Education, Mean (SD)	14.6 (2.6)	14.6 (3.0)	11.3 (4.0)	13.4 (3.0)	
MMSE, Mean (SD)	29.2 (1.1)	29.1 (1.2)	28.0 (1.6)	19.2 (8.4)	
ApoE4+	1 E4	61 (30.5%)	56 (29.2%)	2 (20.0%)	32 (24.6%)
	2 E4	3 (1.5%)	1 (0.5%)	0 (0.0%)	7 (5.4%)
MC = Mutation Carrier; NC = Non-carrier *Table statistics based on 534 participants with confirmed mutation data available as of 6/30/19. Of them 393 (73.6%) are mutation carriers(of these,CDR score is missing for 1), 141 (26.4%) are mutation non-carriers(of these,CDR score is missing for 1),					

Data Sharing Experience (2019)

- 173 data requests granted (82% approval rate)
- 56 tissue requests granted (77% approval rate)
- 149 Publications related to DIAN data

Stages of Proteinopathy in ADAD



Data Request process

The screenshot shows a web browser displaying the 'Data Request Terms and Instructions' page on the DIAN website. The browser's address bar shows the URL: <https://dian.wustl.edu/our-research/observational-study/dian-observational-study-investigator-resources/data-request-terms-and-instructions/>. The website header includes the Washington University School of Medicine in St. Louis logo and the text 'The Dominantly Inherited Alzheimer Network'. A navigation menu lists: HOME, ABOUT, OUR RESEARCH, FOR FAMILIES, FOR HEALTH PROFESSIONALS, NEWS, and DONATE. A search bar is located in the top right of the header. On the left side, there is a vertical menu under 'OUR RESEARCH' with links to: For Investigators, Our Collaborators, Registry, Clinical Trial, Observational Study, Investigator Resources (highlighted), Study Sites, Participation, Procedures, Publications, Leadership, Related Entities, Funding and Study Team, and The Pharma Consortium. The main content area features the title 'Data Request Terms and Instructions' with a sub-link '« back to Investigator Resources'. The text explains the principles of Productivity, Transparency, Fairness, and Inclusiveness, and details the request process. A red button labeled 'SUBMIT A DATA REQUEST' is visible. On the right side, there is a section titled 'Suggested timeline for resource requests' with a list of guidelines. The footer contains contact information for the Dominantly Inherited Alzheimer Network at Washington University School of Medicine, including mailing and office addresses, and social media icons.

Washington University School of Medicine in St. Louis

ENGLISH

The Dominantly Inherited Alzheimer Network

HOME ABOUT OUR RESEARCH FOR FAMILIES FOR HEALTH PROFESSIONALS NEWS DONATE

OUR RESEARCH

For Investigators

Our Collaborators

Registry

Clinical Trial

Observational Study

Investigator Resources

Study Sites

Participation

Procedures

Publications

Leadership

Related Entities

Funding and Study Team

The Pharma Consortium

Data Request Terms and Instructions

[« back to Investigator Resources](#)

We follow the principles of Productivity (with Recognition of the investigator who develops a research idea and does the work to publish it), Transparency, Fairness, and Inclusiveness. The following policies regarding access to DIAN data are intended to provide structure to the request process, respect for intellectual contributions, and standards regarding security/confidentiality.

Request process

All requests for data must be submitted in writing via the electronic data request form available on this website. The request form will include investigator affiliation, contact information, funding support, institutional review board (IRB) approval (if applicable), an NIH-style biosketch, and a brief description of the project, including specific aims, study design, characteristics of the data requested and analysis plans. Investigators requesting data also will be required to sign a data user agreement and an acknowledgement agreement, as specified below. Applications for data use will be reviewed by the Principle Investigator and relevant DIAN Core Leaders. The number, type and disposition of the requests will be tracked by the DIAN Administration Core and a data sharing report will be generated for progress and final reports to the National Institute on Aging (NIA). Each data request will specify the data elements required for the planned analyses. The Biostatistics Core personnel will prepare a file containing only these data elements, together with a participant identification number (not the DIAN ID# but identifier recoded to protect confidentiality) so that questions about particular individuals can be resolved without the investigator's knowledge of the participant's identity. Image data will be available to approved investigators in raw and post-processed formats via the DIAN Central Neuroimaging Data Archive (CNDA) after a formal Data Use Agreement (DUA) has been signed by both institutions.

To respect intellectual contributions of DIAN investigators and collaborators, the following system has been adopted.

- DIAN-affiliated requests: Data analyses to support manuscripts can be developed and proposed by anyone in DIAN: Core Leaders, Site Leaders, other Steering Committee members, or any student, fellow, or investigator designated by a DIAN Site Leader. All data requests will be reviewed by the relevant Core Leaders and the Principal Investigator of DIAN. Upon approval, these requests will receive the most recent data freeze modules.
- External requests: For requests from those not affiliated with the DIAN study, upon approval, access to the previous data freeze will be granted. Such projects/manuscripts will follow the "DIAN-affiliated request" review and approval process (see next section). Publications by investigators outside DIAN would list the DIAN Study Group in the acknowledgments with the required paragraph at the end of this document.

Suggested timeline for resource requests

- Allow 30-60 days for the review process (data only)
- Allow 60-90 days for tissue requests (require external review)
- Allow 30 days for interaction with the Biostatistics Core to provide dataset.

SUBMIT A DATA REQUEST

Dominantly Inherited Alzheimer Network
Washington University School of Medicine
Mailing Address: 660 S. Euclid Ave., CB 8111 | St. Louis, MO 63110
Office Location: 4488 Forest Park Ave. | St. Louis, MO 63108

Contact

Medicine

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EMERGENCY POLICIES NEWS

<https://dian.wustl.edu/our-research/observational-study/dian-observational-study-investigator-resources/>

Comparison of Dominantly Inherited and Sporadic Alzheimer's Disease for regulatory consideration

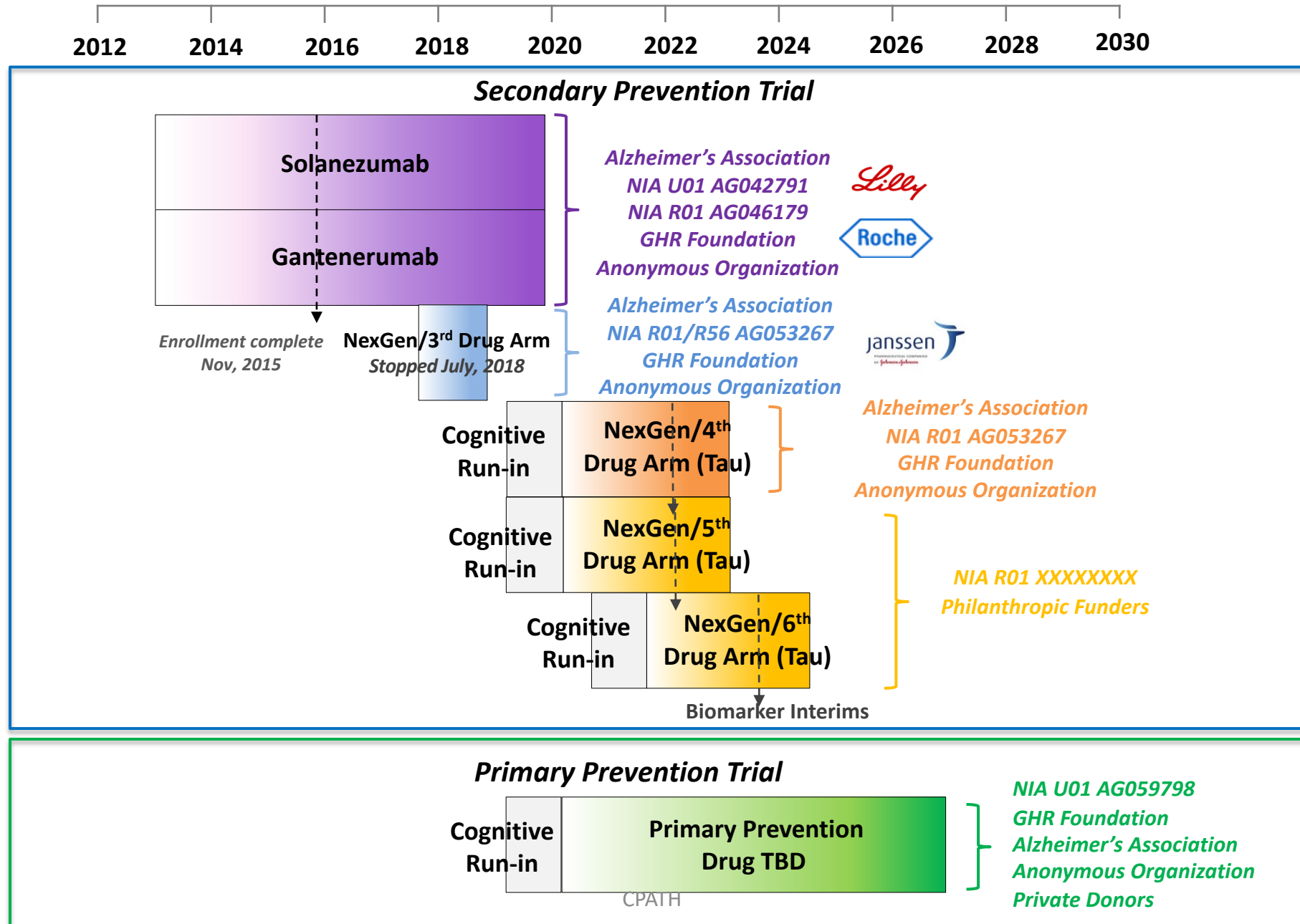
	Dominantly Inherited AD	Sporadic AD
Clinical presentation	Amnestic	Amnestic
Cognitive deterioration	Memory, frontal/executive, generalized cognitive decline	Memory, frontal/executive, generalized cognitive decline
MRI	Hippocampal atrophy and whole brain atrophy	Hippocampal atrophy and whole brain atrophy
PiB PET	Cortex plus basal ganglia	Cortex
FDG PET	Parieto-occipital hypometabolism	Parieto-occipital hypometabolism
CSF A β 42	Decreased by 50%	Decreased by 50%
CSF tau	Increased by 2-fold	Increased by 2-fold

Scientific data supports a Dominantly Inherited AD trial as part of a drug registration package for both Dominantly Inherited and Sporadic AD.

Trials Unit

Maximizing data sharing from prevention trials- DIAN and CAP

DIAN-TU AD Prevention Trial Platform

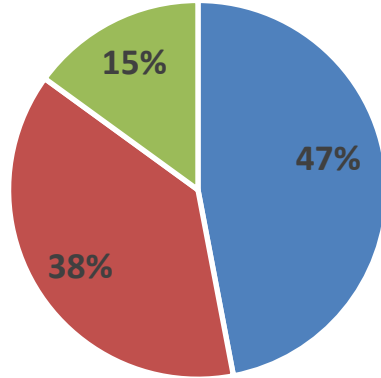




DIAN-TU-Trial: Current Status

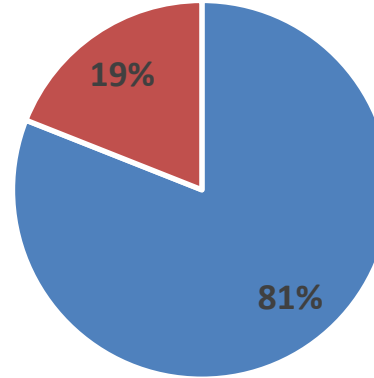


Recruitment Sources



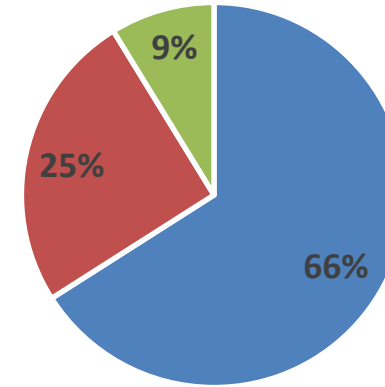
- DIAN Observational Study
- DIAN Expanded Registry
- Sites/Other

Enrollment Metrics



- Randomized
- Screen Fail

Participant Status



- Active
- Early Term - True
- Early Term - Other (mut neg, drop b/f dose)

Test Measure		Longitudinal Assessments per Participant	Quantity	Compliance Rate
Clinical Measures	CDR, CDR-SB, MMSE, FAS, GDS, NPIQ	5	≈1,000	100%
Cognitive Measures	Cogstate, Pencil / Paper	10-15	≈ 1,525	98%
Fluid Biomarkers	Plasma, Serum, CSF	4	≈ 1,450	99%
Imaging Biomarkers	MRI, PiB, AV-45, FDG, AV-1451	4	≈ 5,000	99%

Collaboration for Alzheimer's Prevention (CAP)

- Collaboration and coordination of prevention trial efforts
- Alzheimer's Association, National Institute of Aging, Fidelity Foundation, FDA, **API, ADCS A4, and DIAN-TU**
- Harmonization of biomarkers, clinical, and cognitive measures, and sharing of approaches for prevention efforts.
- Data and sample sharing - with NIA funding, data and samples will be made available to qualified investigators per an approved resource sharing plan.

DIAN-TU's data and sample sharing principles:

- Protection of trial integrity
- Protection of participants
(confidentiality of mutation status)
- Scientific Advancement
- Productivity
- Transparency
- Fairness
- Inclusiveness

Publication benefits to DIAN-TU trial Cores and investigators

- **Baseline Data:** Cores, site investigators, and trial pharma partners have the opportunity to publish in the **first year** (after enrollment is complete and data is frozen)
- **Post-Randomization Data:** Cores, site investigators, and trial pharma partners have the opportunity to publish in the **first 18 months** after drugs arms complete all necessary regulatory work
- After these periods, data is available to qualified investigators

Resources

Websites:

- DIAN & DIAN-TU: <https://dian.wustl.edu/>
- DIAN Expanded Registry: <https://dian.wustl.edu/our-research/registry/>

Contact Information:

- DIAN EXR email: dianexr@wustl.edu
- DIAN EXR Coordinator: **844-DIAN-EXR (844-342-6397)**
- DIAN Observational Deputy Director **314-747-1940**

The DIAN (NIH UF1AG032438)

The DIAN participants and family members

The Alzheimer's Association, ADAD Forum, DIAN Pharma Consortium



Argentina

Fundación para la Lucha contra las Enfermedades Neurológicas de la Infancia (FLENI), *Ricardo Allegri*

Australia

Mental Health Research Institute, Melbourne, *Colin Masters*

Edith Cowan Univ , Perth, *Ralph Martins*

Prince of Wales Medical Research Institutes, Sydney, *Peter Schofield*

Germany

Ludwig-Maximilians-Universität München, *Adrian Danek*

University of Tübingen, *Matthias Jucker*

Japan: DIAN-Japan, Mori

Hirosaki University, *Mikio Shoji*

Niigata University, *Takeshi Ikeuchi*

Osaka City University, *Hiroyuki Shimada*

University of Tokyo, *Kazushi Suzuki*

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United Kingdom

Institute of Neurology, Univ College London, *Martin Rossor*

United States

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University of Pittsburgh, *William Klunk*

Butler Hospital/Brown University, *Steve Salloway*

DIAN-TU Administrative and Clinical Operations Team

Randall Bateman – Director and PI, Eric McDade – Associate Director

Sarah Adams, Stephanie Belyew, Erin Cattoor, David Clifford, Garland Edmonds, Molly Fitzgerald, Erica Fowler, Amanda Fulbright, Angela Fuqua, Kurtis Hanks, Ron Hawley, Dottie Heller, Latoya Jones, Michelle Jorke, Nicole Kelley, Paulette MacDougall, Jacki Mallmann, Tayona Mayhew, Karen McCann, Susan Mills, Julie Murphy, Jennifer Petranek, Anna Santacruz, Jessi Smith, Annette Stiebel, Shannon Sweeney, Linda Watkins-Imhof, Ellen Ziegemeier

DIAN-TU Cores

Administrative: Randall Bateman, Eric McDade and team

Biomarkers: Anne Fagan and team

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Cognition: Jason Hassenstab and team

Genetics: Alison Goate, Carlos Cruchaga and team

Imaging: Tammie Benzinger and team

Neuropathology: Rick Perrin and team

DIAN-TU Collaborators

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Granzer Regulatory Consulting , IQVIA

DIAN-TU Therapy Evaluation Committee: Paul Aisen, Randall Bateman,

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Jeffrey Kelly, Virginia Lee, Cynthia Lemere, Eric McDade, Erik Musiek,

Erik Roberson, Matthias Staufenbiel, Robert Vassar

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David Knopman, Kristine Yaffe

ADCS: Ron Thomas **ATRI:** Paul Aisen

University of Michigan: Robert Koeppe

Mayo Clinic: Clifford Jack

We gratefully acknowledge the DIAN and DIAN-TU participants and family members, DIAN Team, DIAN Steering Committee, Knight ADRC, Alzheimer's Association, ADAD Forum, NIH U01AG042791, R01AG046179, R01/R56AG053267, U01AG059798, R13AG055232, DIAN-TU Pharma Consortium, GHR, Anonymous Foundation, Industry Partners (Eli Lilly & Co., Hoffman-LaRoche, Janssen, Avid Radiopharmaceuticals, Bracket, Cogstate), and Regulatory Representatives.

DIAN-TU Sites

Australia

Mental Health Research Institute, *Colin Masters*
McCusker Foundation for AD Research, *Roger Clarnette*
Neuroscience Research Australia, *William Brooks*

Canada

University of British Columbia, *Robin Hsiung*
Sunnybrook Health Sci Centre, *Mario Masellis*
McGill University, *Serge Gauthier*
CHU de Quebec - Hôpital de l' Enfant Jésus, *Robert Laforce**

France

Hôpital Roger Salengro, *Florence Pasquier*
Hôpital Neurologique Pierre Wertheimer, *Maité Formaglio*
Groupe Hospitalier Pitie-Salpetriere, *Bruno Dubois*
Hôpital Charles Nicolle, *David Wallon*
Hôpital Purpan, *Jérémie Pariente*

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The National Hospital for Neurology & Neurosurgery,
Catherine Mummery

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Yale University, *Christopher van Dyck*
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University of Pittsburgh, *Sarah Berman*
University of Puerto Rico, *Ivonne Jimenez-Velazques*
Butler Hospital/Brown University, *Ghulam Surti*
University of Texas Southwestern Medical Center, *Diana Kerwin**
University of Washington, Seattle, *Suman Jayadev*

*New sites pending approval