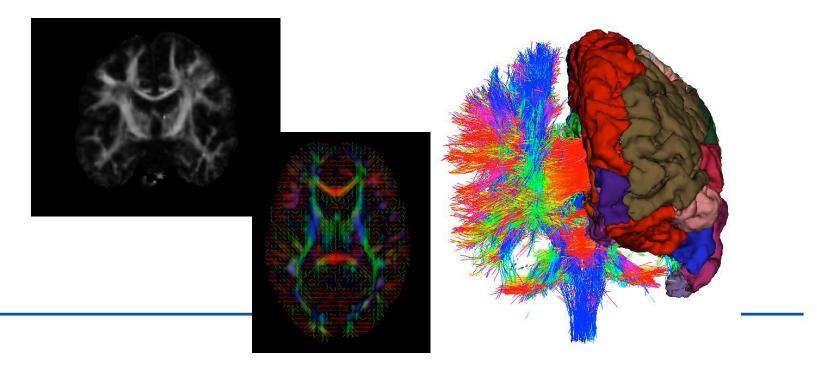


# NA-MIC National Alliance for Medical Image Computing http://www.na-mic.org

# Driving Biological Problem Huntington's Disease THE UNIVERSITY OF IOWA

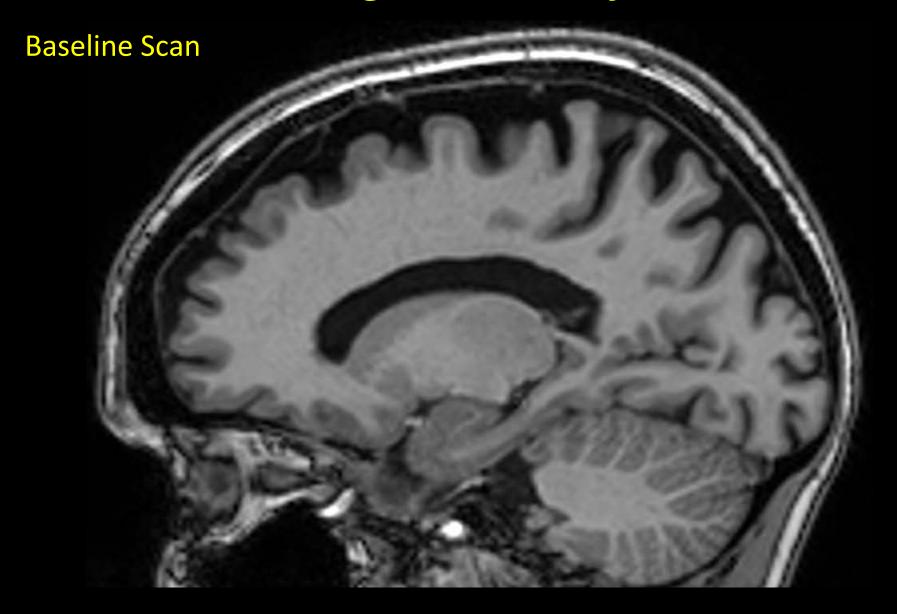




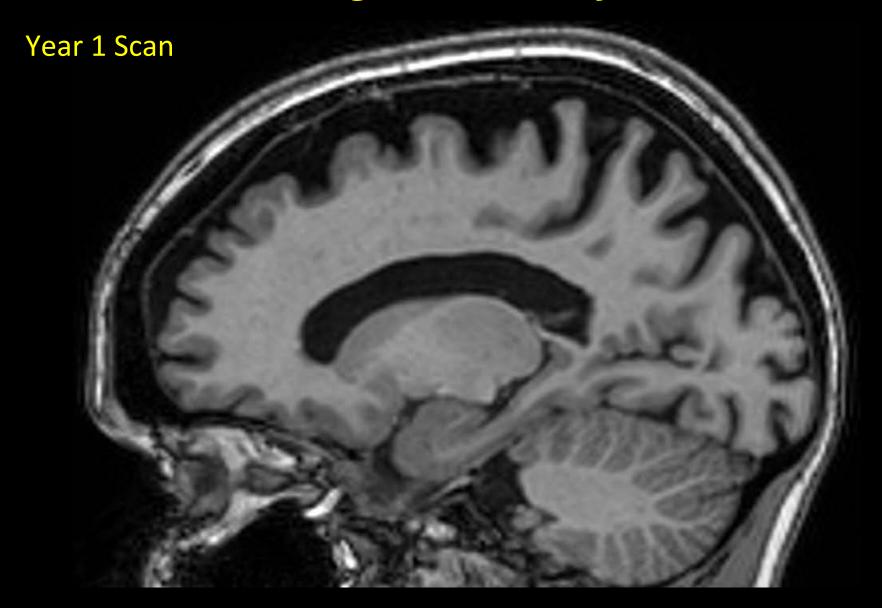
### **Builds on PREDICT-HD**

 The NIH-funded project "Neurobiological Predictors of Huntington's Disease" (PREDICT-HD) studies Huntington's disease (HD), a neurodegenerative genetic disorder that affects muscle coordination, behavior, and cognitive function, and causes severe debilitating symptoms by middle age.

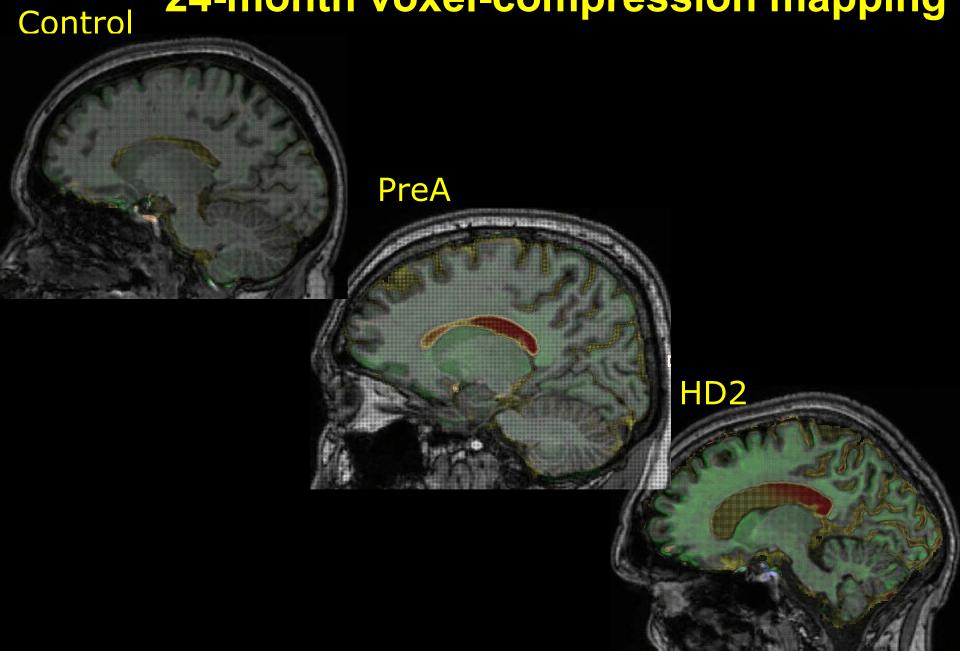
# **TRACK-HD Stage 1 HD Subject**



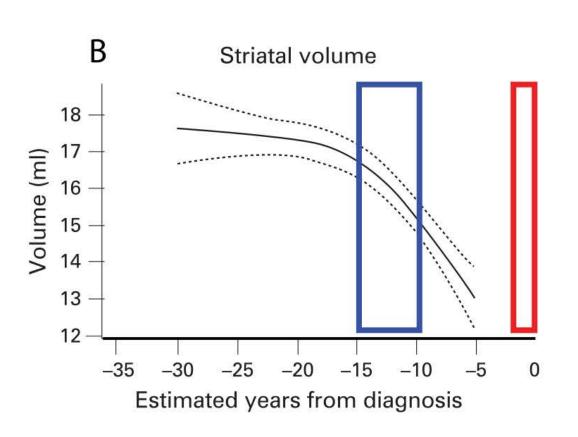
# **TRACK-HD Stage 1 HD Subject**

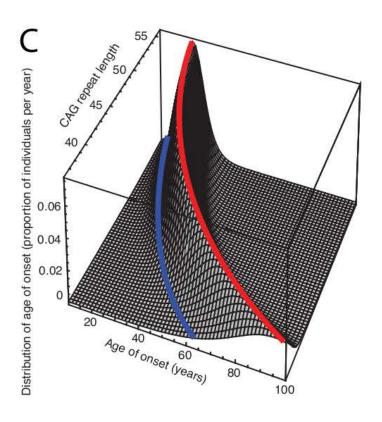


### 24-month voxel-compression mapping



## **HD Background**







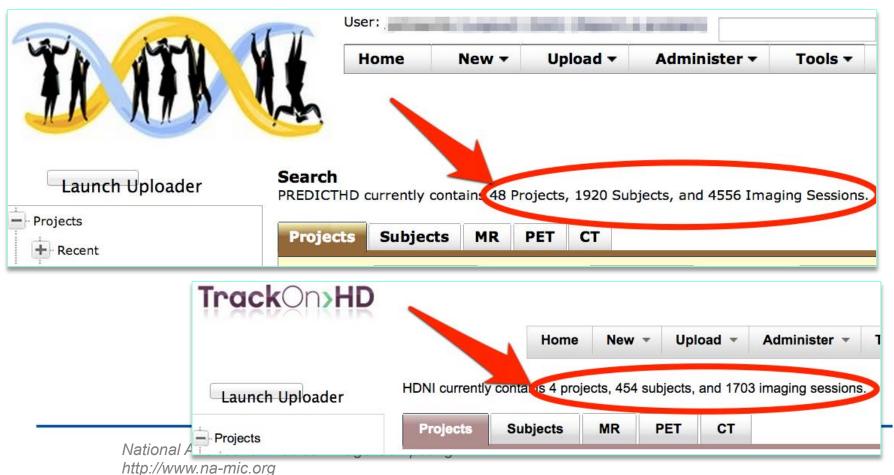
## **Specific Aims**

- Perform individualized longitudinal shape change quantification from multi-modal data.
- Complete full brain Diffusion Tensor Imaging tractography analysis.
- Deploy extensible tools for sharing source data, derived data, algorithms and methods to multi-site analysis teams.



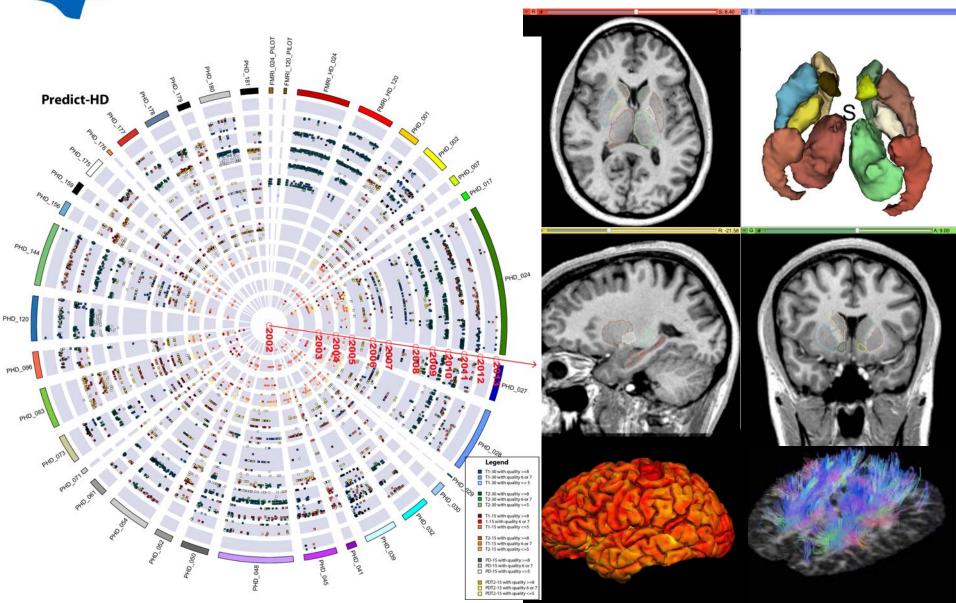
# **Sharing HD Data**

http://www.na-mic.org/Wiki/index.php/2011\_Summer\_Project\_Week



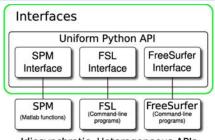


# **Improve Data Processing**

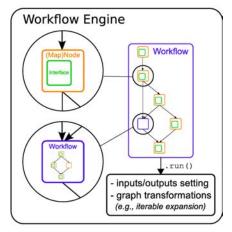


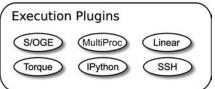


# NiPype: Large catalog of tools with a uniform interface



Idiosynchratic, Heterogeneous APIs





#### Batch processing

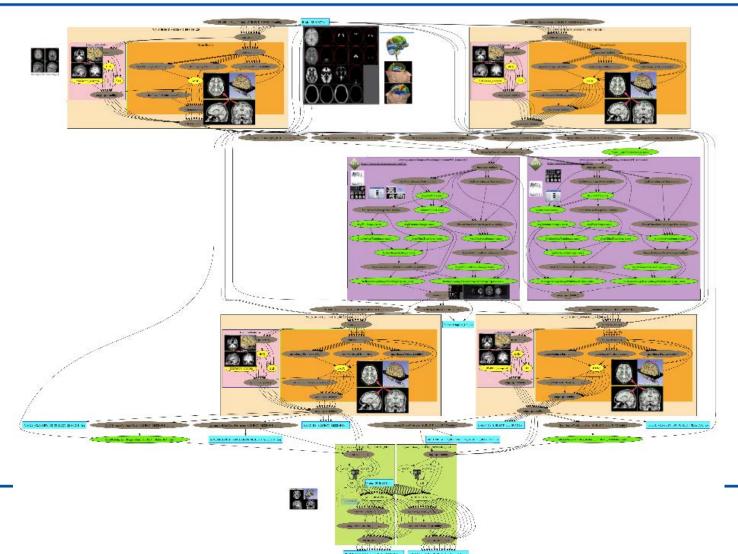
- Distributed processing plugins
- Reruns affect updated/edited node connections ONLY!
- Uniform node creation
  - Stable and consistent API
  - Nipype's Function node allows easy integration of CLI tools

### Pipeline complexity

- Iterables, MapNodes
- Nested workflows

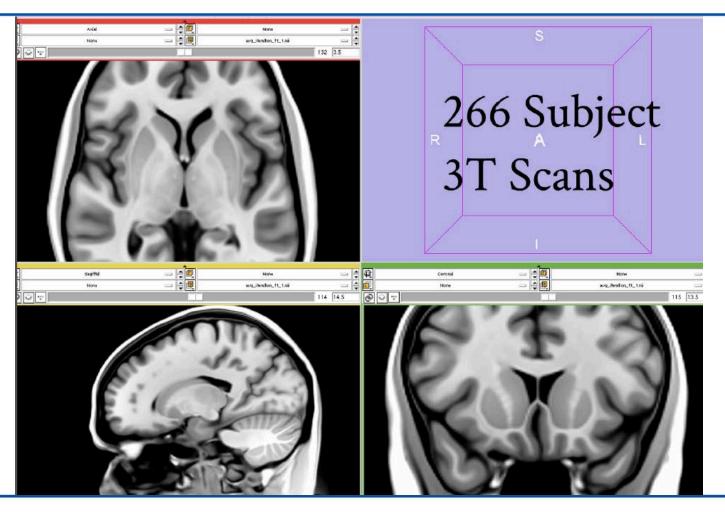


# Diagram of Longitudinal Processing Pipeline





# Improved Template Building with ANTS



National Alliance for Medical Image Computing http://www.na-mic.org



# **Brain Sub-Cortical Structures:**BRAINSCut (Longitudinal estimation)

#### Developed

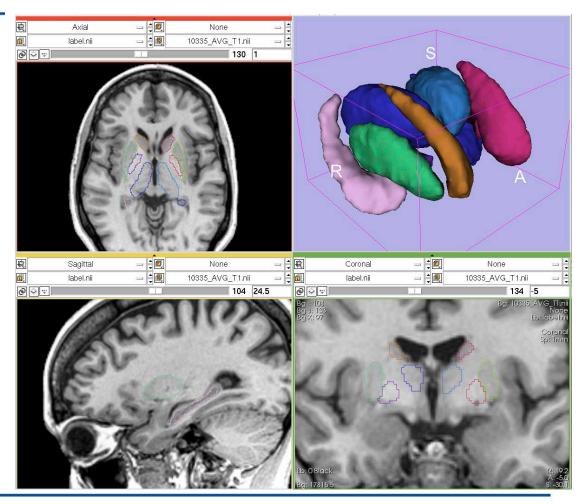
- Caudate
- Putamen
- Thalamus

#### New structure

Hippocampus

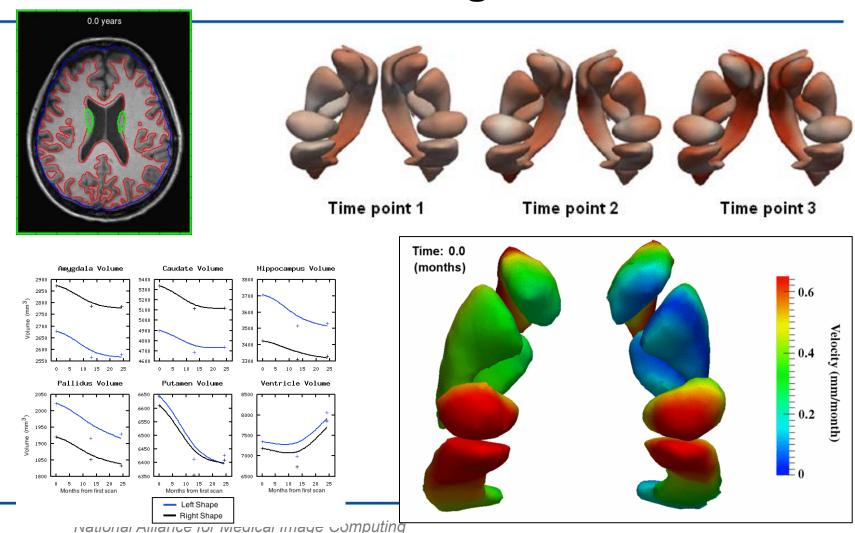
### On going

- Globus
- Accumbens
- And more...





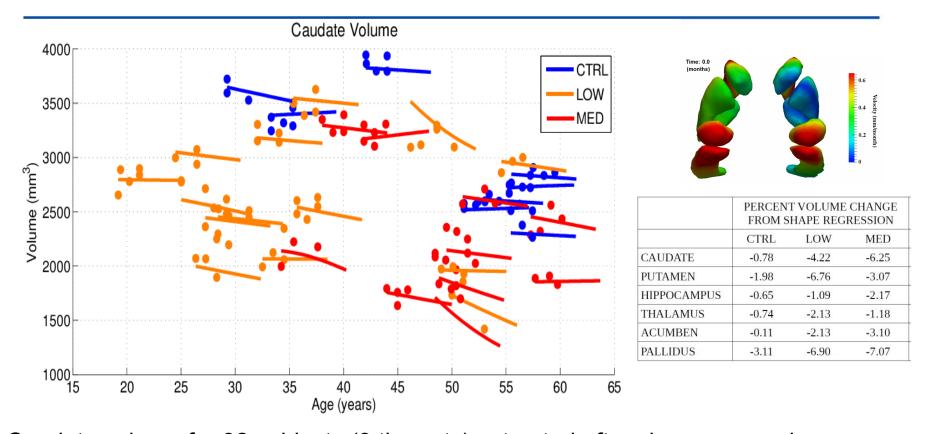
# HD: Joint 4D modeling of subcortical structures from longitudinal MRI



http://www.na-mic.org



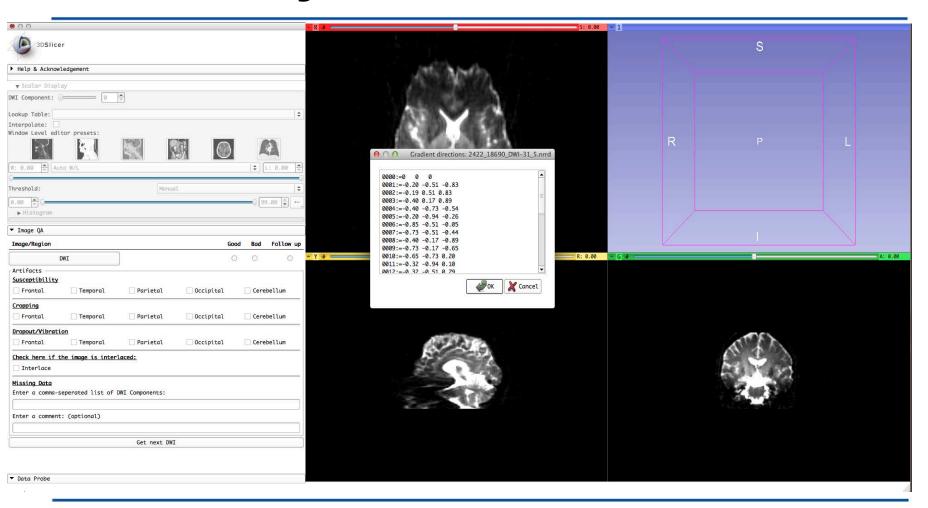
# Personalized Shape Trajectory Modeling across Subjects



Caudate volume for 32 subjects (3 time pts) extracted after shape regression. Observed volumes are shown as circles, which highlight the noise in segmentation.

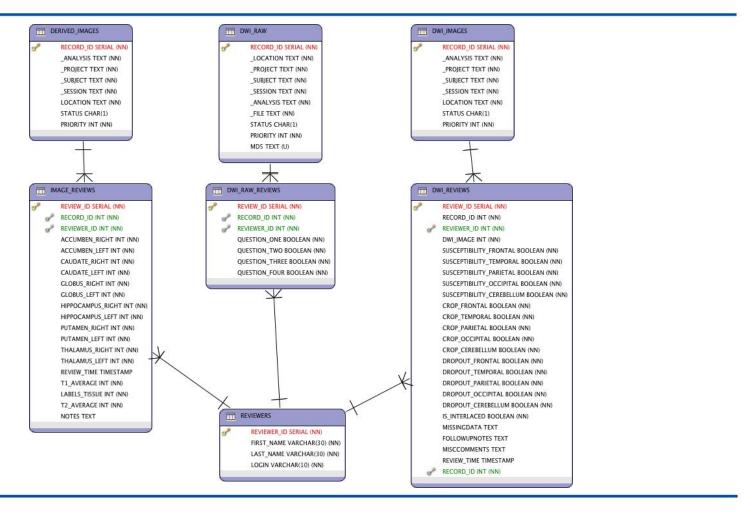


## **Quality Assurance tools**





## **Quality Assurance tools**





# **Data Reports**

A	В		С		D	E	F	
project	subject	v	session	•	ICV_2088: ▼	l_caudate_	r_caudate_1	
HDNI_001	015955315		015955315_20100803_	30	1494658	829		
HDNI_001	015955315		015955315_20090622_	30	1498021	947	1	
HDNI_001	045416487		045416487_20110614_	30	1413108	1714	1	
HDNI_001	045416487		045416487_20080522_	30	1412544	1941	21	
HDNI_001	068044003		068044003_20090521_	30	1770211	2580	2.	
HDNI_001	068044003		068044003_20100628_	30	1756838	2575	2	
HDNI_001	068044003		068044003_20080609	30	1746901	2729	2	

A	В	C	D	E	F	
"column_descriptors_id"	"column_descriptors_description	"measurement_code_label"	"biological_region_label"	"experiment_label"	"tool_label"	
cranial_csf_358		mm_3	cranial_csf	20130109_TrackOn_Results	BRAINSTools_20130109	
r_hippocampus_162		mm_3	r_hippocampus	20130109_TrackOn_Results	BRAINSTools_20130109	
ICV_20882		mm_3	ICV	20130109_TrackOn_Results	binaryBrainMask_20130401	
r_caudate_134		mm_3	r_caudate	20130109_TrackOn_Results	BRAINSTools_20130109	
_thalamus_106		mm_3	I_thalamus	20130109_TrackOn_Results	BRAINSTools_20130109	
globus_330		mm_3	I_globus	20130109_TrackOn_Results	BRAINSTools_20130109	
cerebellar_white_matter_20938 /		mm_3	cerebellar_white_matter	20130109_TrackOn_Results	BRAINSTools_20130109	
datase	data_dictionary comments_scores	(+)	****		10. VARV	

A	В	С	D	E	
project_label	subject_label	session_label	column_descriptors_id	scores	comm
HDNI_001	_001 560216908		I_hippocampus_50	75	Visual in
IDNI_003 034032275		034032275_20121001_30	r_putamen_190	95	Visual in
HDNI_002	058773254	058773254_20080609_30	I_putamen_78	95	Visual in
HDNI_003	658092030	658092030_20080204_30	r_globus_302	95	Visual in
HDNI_003	867249739	867249739_20110427_30	I_thalamus_106	95	Visual in
HDNI_003	541359481	541359481_20080611_30	r_caudate_134	95	Visual in
HDNI_004	287905458	87905458_20120801_30	r_thalamus_218	95	Visual in
	set data_dictionar comments	scores			9

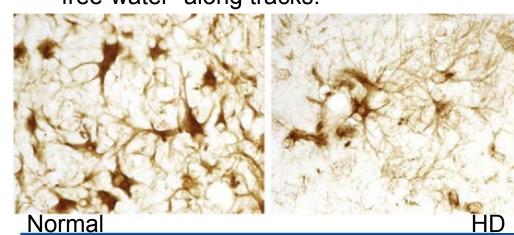
http://www.na-mic.org

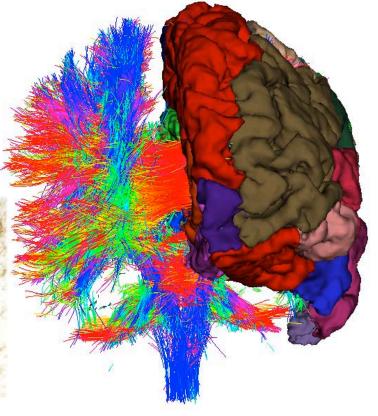


## **Tractography**

 It is clear that a more comprehensive approach is needed that could easily be extended as new questions arose. (i.e. support more exploration than direct apriori interrogation)

 Van Camp 2012 Showed that loss of neurons can be measured by estimating "free-water" along tracks.





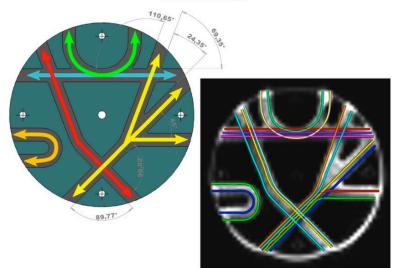


# What is best tractography?

Quantitative evaluation of 10 tractography algorithms on a realistic diffusion MR phantom

Pierre Fillard a,b,\*, Maxime Descoteaux c, Alvina Goh d, Sylvain Gouttard e, Ben Jeurissen f, James Malcolm g, Alonso Ramirez-Manzanares h, Marco Reisert i, Ken Sakaie j, Fatima Tensaouti k, Ting Yo i,

Jean-François Mangin b, Cyril Poupon m



a Parietal Research Team, INRIA Saclay Île-de-France, Neurospin, France

b Laboratory of Computer-Assisted Neuro-Imaging, CEA Saclay, Neurospin, France

<sup>&</sup>lt;sup>c</sup> MOIVRE Center, Computer Science Department, Université de Sherbrooke, Canada

<sup>&</sup>lt;sup>d</sup> Department of Mathematics, National University of Singapore, Singapore

e Scientific Computing and Imaging Institute, University of Utah, USA

f IBBT-VisionLab, Department of Physics, University of Antwerp, Belgium

g Psychiatry Neuroimaging Laboratory, Brigham and Womens Hospital, Harvard Medical School, USA

h Mathematics Department, University of Guanajuato, Mexico

Department of Radiology, Medical Physics, University Hospital Freiburg, Germany

j Imaging Institute, The Cleveland Clinic, Cleveland, USA

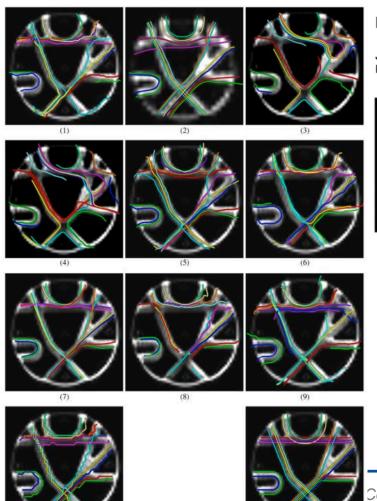
k National Institute for Medical Research, INSERM, U825, France

<sup>1</sup> Max Planck Institute for Human Cognitive and Brain Sciences, Germany

m Imaging and Spectroscopy Laboratory, CEA Saclay, Neurospin, France

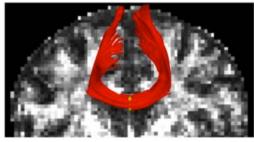


# From Publication To Clinical Application



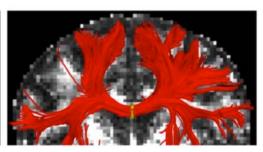
#### Filtered multi-tensor tractography

James G. Malcolm, Martha E. Shenton, and Yogesh Rathi
Psychiatry Neuroimaging Laboratory, Brigham and Womens Hospital, Harvard Medical School

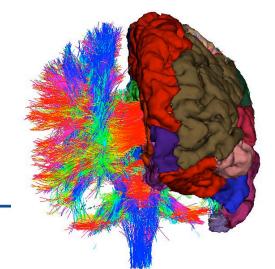


(a) Single-tensor streamline

\* We Choose the Unscented Kahlman Filter algorithm "UKFTractography" \* Improved speed and integration with Slicer



(b) Filtered two-tensor



Computing



# How to select tracts of interest?

### WMQL – White Matter Query Language

Table 1: Association Tract Definitions in WMQL: Cingulum bundle (CG); Extreme Capsule (EmC); and the fascicules: Superior Longitudinal (SLF) sections I to III; Arcuate (AF); Inferior occipito frontal (IOFF); Middle Longitudinal (MdLF); Uncinate (UF)

CB.side = only((cingular.side or cingular\_cortex.side) and (middle\_frontal.side or cuneus.side or entorhinal.side or superior\_frontal.side or inferior\_parietal.side or fusiform.side or medial\_orbitofrontal.side or lateral\_orbitofrontal.side or parahippocampal.side or precuneus.side or lingual.side or centrum\_semiovale.side))

EmC.side = (endpoints\_in(inferior\_frontal.side or middle\_frontal.side)
 andendpoints\_in(inferior\_parietal\_lobule.side) andtemporal.side and insula.side)not in
hemisphere.opposite

SLF\_I.side = (superior\_parietal.side and precuneus.side and superior\_frontal.side)

or(superior\_parietal.side and precuneus.side and superior\_frontal.side and
lateral\_occipital.side) not in cingular.side not in temporal.side not in subcortical.side not in
hemisphere.opposite

SLF\_II.side =(inferior\_parietal.side or supramarginal.side or lateral\_occipital.side) and endpoints\_in(middle\_frontal.side)) not in temporal.side not in subcortical.side not in hemisphere.opposite

SLF\_III.side = ((inferior\_parietal.side or supramarginal.side or lateral\_occipital.side) and endpoints\_in(inferior\_frontal.side)) not in temporal.side not in subcortical.side not in hemisphere.opposite

AF.side = (inferior\_frontal.side or middle\_frontal.side or precentral.side) and
(superior\_temporal.side or middle\_temporal.side) not in medial\_of(supramarginal.side)not in
subcortical.side not in hemisphere.opposite

IOFF.side = (lateralorbitofrontal.side and occipital.side) and temporal.side not in subcortical.side not in cingular.side not in superior\_parietal\_lobule.side not in hemisphere.opposite

ILF.side = only(temporal.side and occipital.side) and anterior\_of(hippocampus.side) not in

Anatomical Terms



b) endpoints\_in(postcentral)



c) postcentral not in only(postcentral) not in endpoints\_in(postcentral)

Relative Position Terms



d) superior\_of(amygdala) inferior\_of(amygdala)



e) anterior\_of(amygdala) posterior\_of(amygdala)



f) lateral\_of(amygdala) medial\_of(amygdala)







g) a = insula and h (inferior\_frontal or middle\_frontal or orbito\_frontal)





b = temporal and anterior\_of(amygdala)



i) uncinate\_fasciculus = a and endpoints\_in(b)



# Dispersion is an important measure

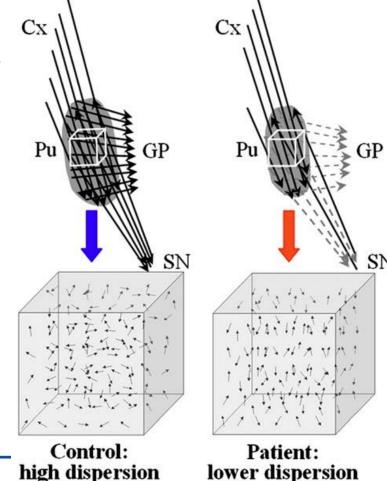
 [Douaud09] describes the selective loss in fibers leading to lower dispersion in HD subjects (left).

Need to translate methods that enable measurement of

**dispersion:** A fiber tract connecting the substantia nigra in the brainstem to the striatum is colored by fiber

dispersion [Savadjiev12].

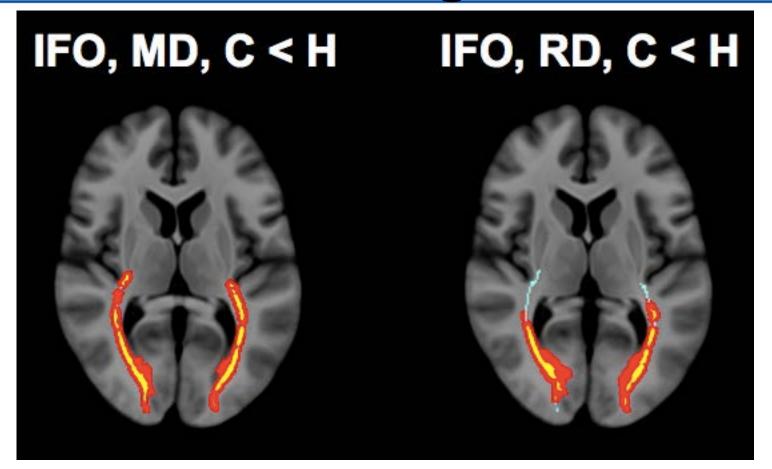
Working with Peter
Savadjiev & Yogesh Rathi
to transfer this
methodology from
prototype environment to
production environment



National Alliance for Medical Image Computing http://www.na-mic.org



# Preliminary Longitudinal DWI Processing Results





Laurent Younes, J. Tilak Ratnanather, Timothy Brown, Elizabeth Jane S. Paulsen, Russell L. Margolis, Roger L. Albin, Michael I. Miller, and the PREDICT-D Investigators and Coordinators

Jane S. Paulsen, \*\* Russell L. Margolis, \* Roger L. Albin, \* Michael I. Miller, \* and the PREDICTD Investigators and Coordinators

#### frontiers in **NEUROINFORMATICS**

Huntington's Disease is Associated with Regionally

Specific Atrophy

Rachael I. Scahill, 18 Nicola Z. Hobbs, Miranda J. Say, Natalie Bechtel, 2 Susie M.D. Henley, Harpreet Hyare, Douglas R. Langbehn, Rebecca Jones, Blair R. Leavitt, Raymund A.C. Roos, Alexandra Durr, Hans Johnson, Stephane Lehéricy, David Craufurd, Christopher Kennard, Stephen L. Hicks, Julie C. Stout, Ralf Reilmann, Sarah J. Tabrizi

and the TRACK-HD investigators





### UNC-Utah NA-MIC framework for DTI fiber tract analysis

Audrey R. Verde<sup>1\*</sup>, Francois Budin<sup>1</sup>, Jean-Baptiste Berger<sup>1</sup>, Aditya Gupta<sup>1,2</sup>, Mahshid Farzinfar<sup>1</sup>, Adrien Kaiser<sup>1</sup>, Mihye Ahn<sup>3</sup>, Hans Johnson<sup>4</sup>, Joy Matsui<sup>4</sup>, Heather C. Hazlett<sup>1</sup>, Anuja Sharma<sup>5</sup>, Casey Goodlett<sup>6</sup>, Yundi Shi<sup>1</sup>, Sylvain Gouttard<sup>5</sup>, Clement Vachet <sup>1,5</sup>, Joseph Piven<sup>1</sup>, Hongtu Zhu<sup>3</sup>, Guido Gerig<sup>5</sup> and Martin Styner 1,7



E.Y. Regina Kim<sup>1,2</sup>, Hans J. Johnson<sup>1,2</sup> and Norman K. Williams<sup>1</sup> Derivation of fiber tracts representing the corticospinal tract using anatomical landma

Joy Matsui<sup>1,2</sup>, Eun Young Kim<sup>1</sup>, Vincent Magnotta<sup>3,1</sup>, and Hans Joh <sup>1</sup> University of Iowa, Department of Psychiatry, Iowa City, IA, US, - University of Iowa, Department of Psychiatry, Iowa City, III, O. 2. University of Hawaii, John A. Burns School of Medicine, Honolulu, H. Inversity of Hawaii, John A. Daries Believe of Radiology, Iowa City, IA, US

3 University of Iowa, Department of Radiology, Iowa City, IA, US

Segmentation of Multi-me 25October 2013 ificial Neural Netw

g Kim<sup>a,b,\*</sup> and Hans Johnson

402 Seamans Center for the E of Iowa, Iowa city, IA, USA 5 awkins Drive, W278 GH, Iowa

Elizabeth H Aylward, Peggy C Nopoulos, Christopher A Ross, Douglas R Langbehn, 2 Ronald K Pierson, <sup>2</sup> James A Mills, <sup>2</sup> Hans J Johnson, <sup>2</sup> Vincent A Magnotta, <sup>2</sup> Andrew R Juhl, Jane S Paulsen, the PREDICT-HD Investigators and Com-

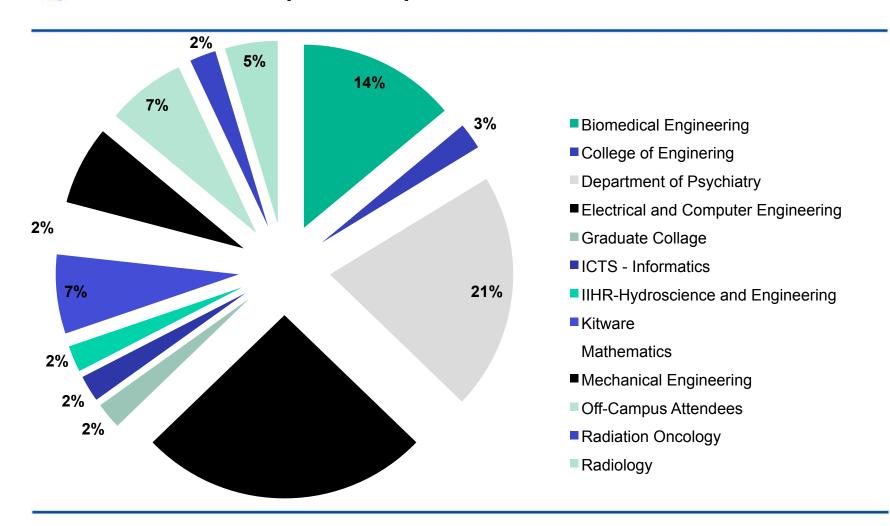
Biological and clinical changes in premanifest and early Huntington's disease in the TRACK-HD study:

And Many Under



### ITK/Simple ITK

#### **ITK/Simple ITK Departmental Attendance**



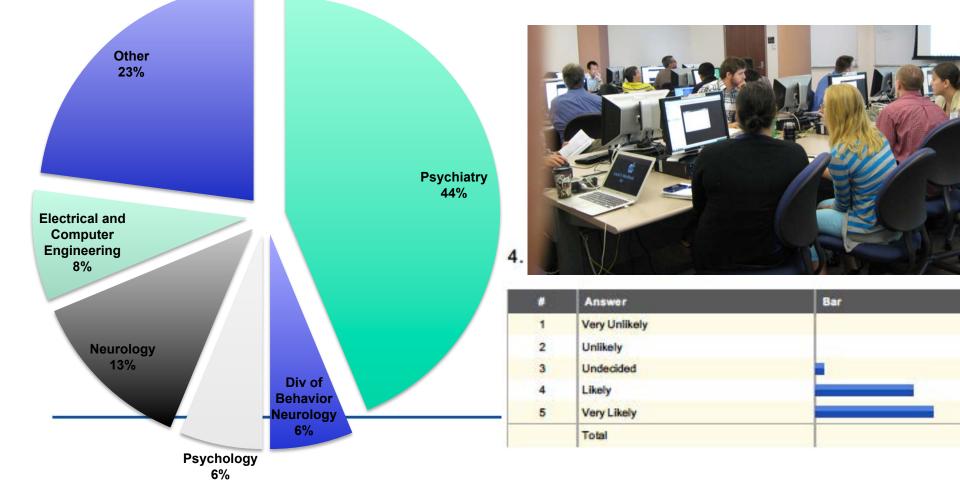


### **BRAINSCamp 2013**

NA-MIC

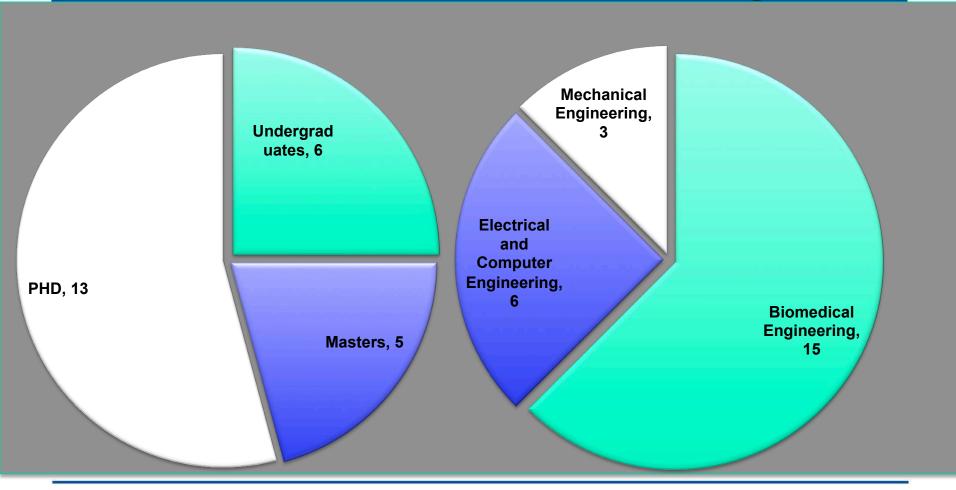
National Alliance for Medical Image Computing http://www.na-mic.org

### % of Departmental Attendance 41 Attendees





### Medical Image Processing Course 24 New "NA-MIC Certified" Engineers





## **Specific Aims**

- Perform individualized longitudinal shape change quantification from multi-modal data. (DONE!)
- Complete full brain Diffusion Tensor Imaging tractography analysis. (DONE!)
- Deploy extensible tools for sharing source data, derived data, algorithms and methods to multi-site analysis teams. (DONE!)



# Funded HD-Projects Using NAMIC DWI Processing Tools

- (CHDI) TRACK-HD/TRACK-ON (Private Foundation)
- (R01 NS040068 12S1) PREDICT-HD
  - (U01 NS082083) Functional Connectivity in Pre-manifest HD (Stephen Rao, Cleveland Clinic)
  - (U01 NS082085) Study of Basal Ganglia Shape Analysis and Circuitry (Michael Miller & Chris Ross, JHU)
  - (U01 NS082086) 4D Shape Analysis for Modeling Spatiotemporal Change Trajectories in Huntington's (Guido Gehrig, Utah)
  - (U01 NS083223) Characterization of White Matter in HD Using Diffusion MRI (CF Westin, Harvard)
  - (U01 NS082074) Imaging and Genetics in Huntington's Disease (Jessica Turner & Vincent Calhoun, MIND Institute)
- (U54 EB005149) DBP-National Alliance For Medical Image Computing



## **Thank You!**

Questions?