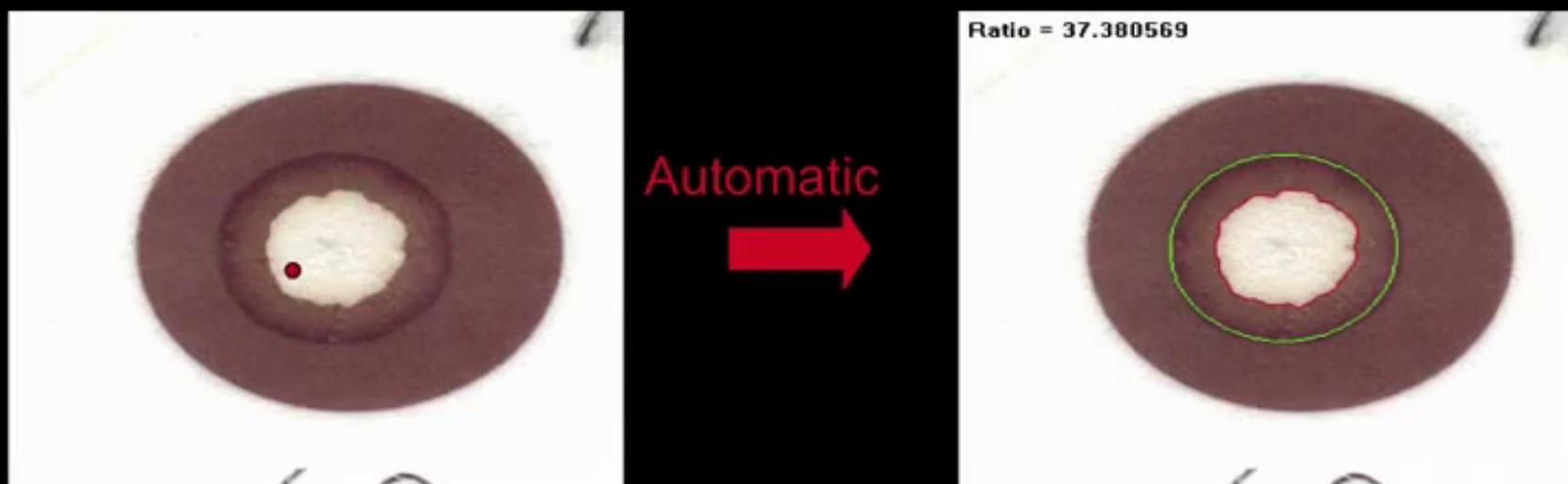


Automatic skin lesion segmentation



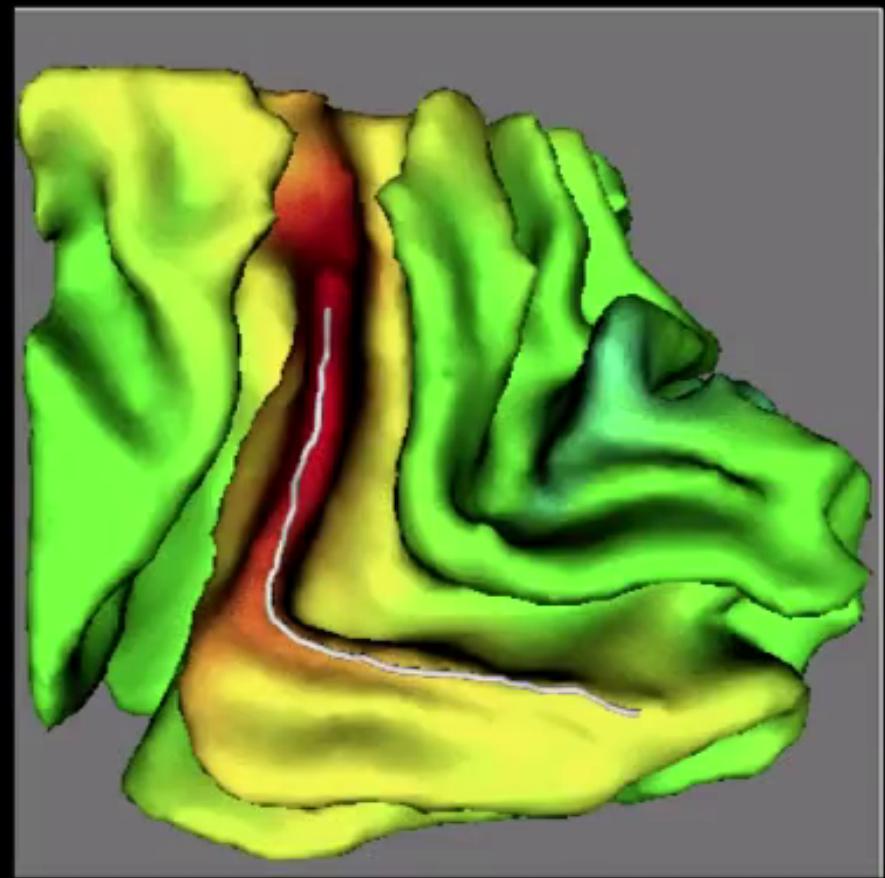
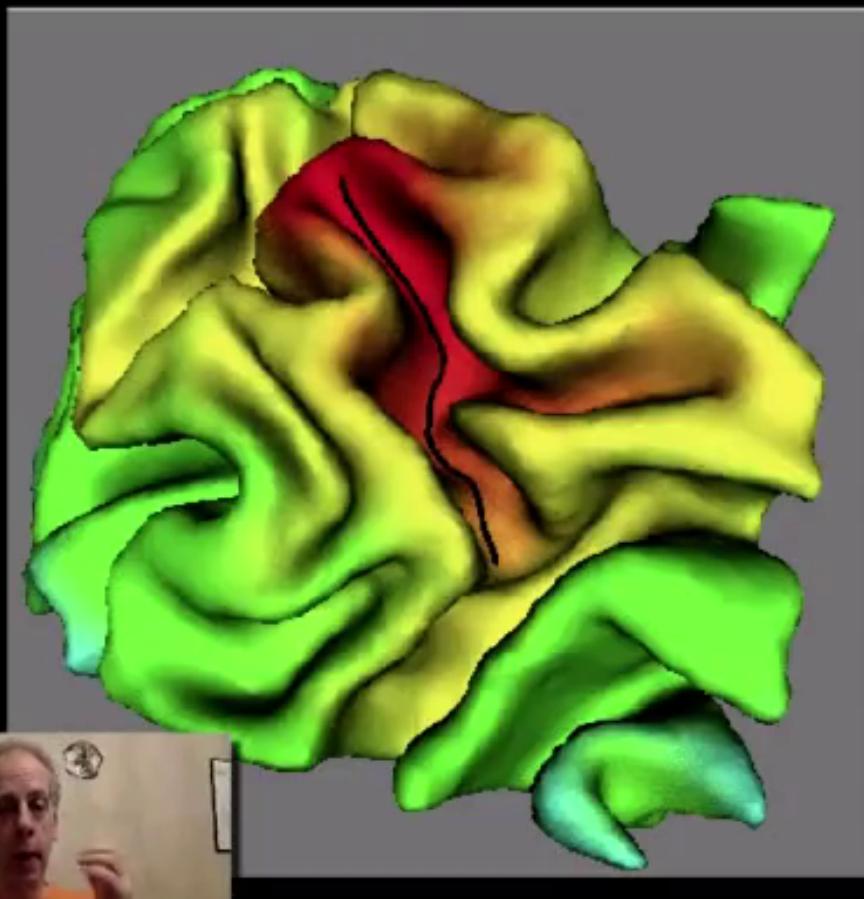
A non-invasive test to aid in the diagnosis of cystic fibrosis: Automatic chloride patch/sensor analysis



- Ratio between red and green areas is in correlation with chloride concentration, aiding in the diagnosis of CF
- Courtesy of PolyChrome and Warren Warwick



Sulcii extraction



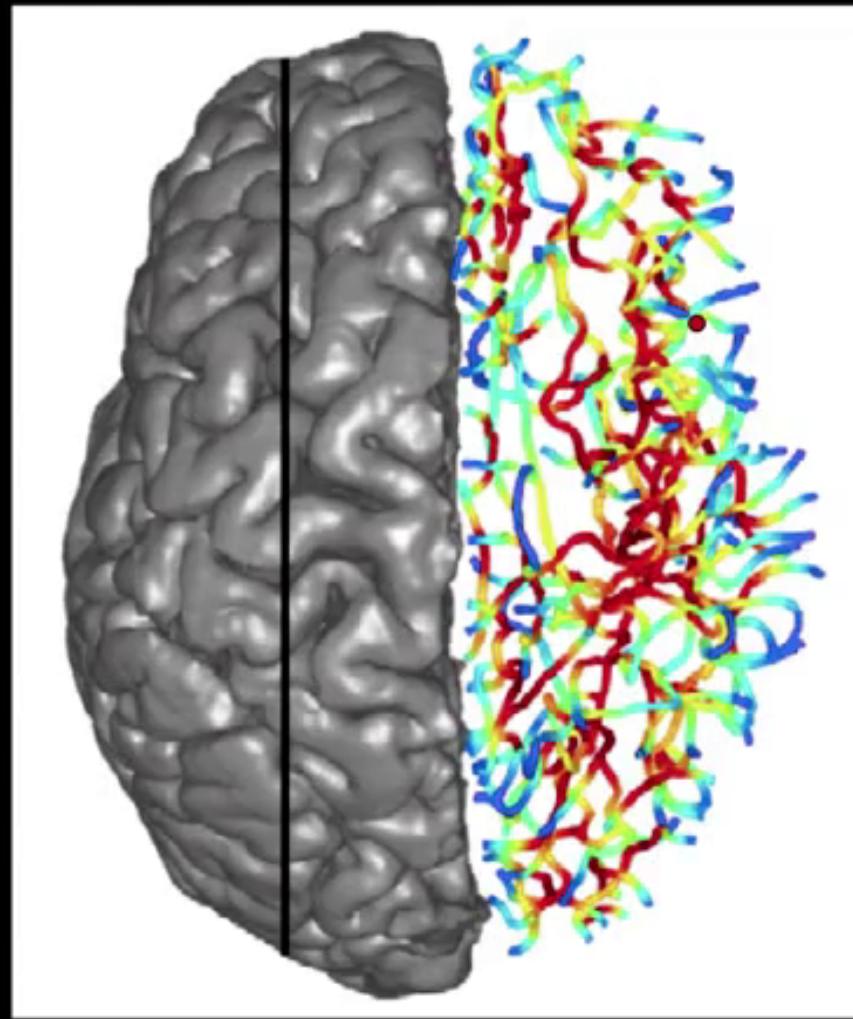


Image Registration, Classification and Averaging in Cryo-Electron Tomography

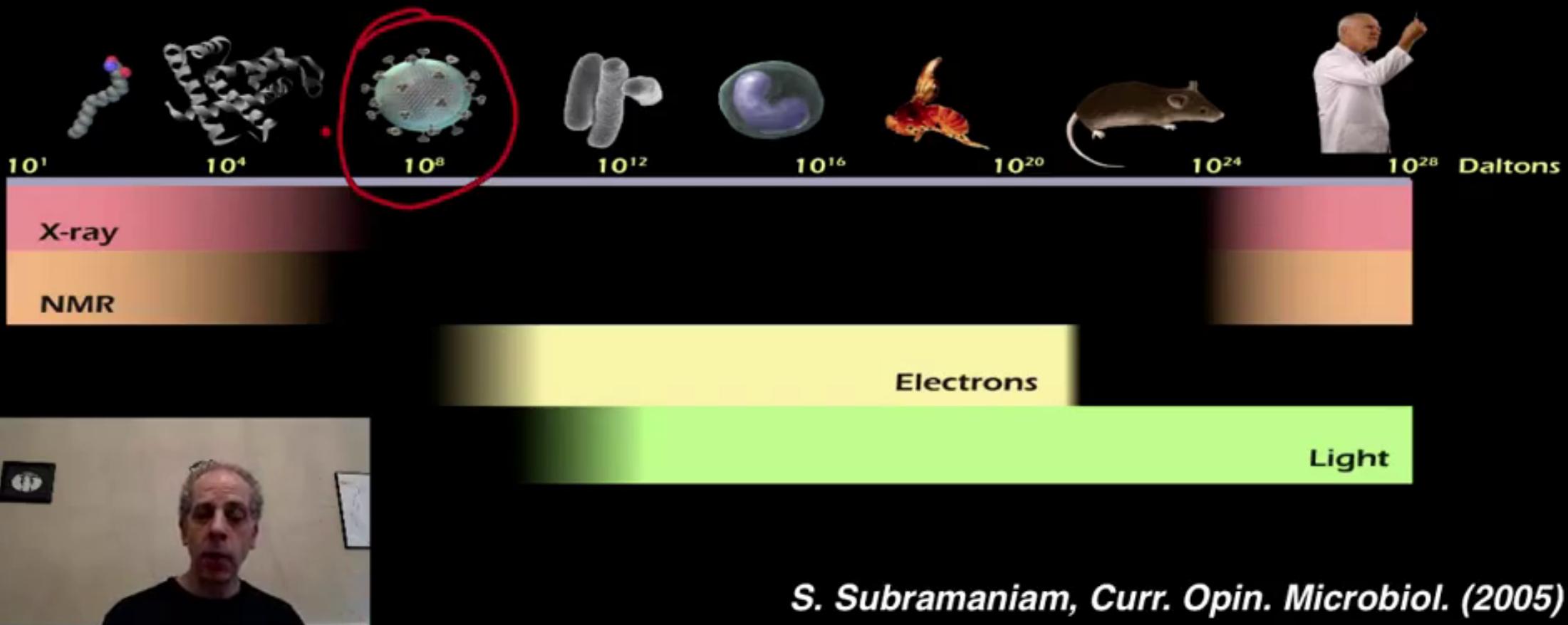
Thanks to Alberto Bartesaghi and Sriram Subramaniam

Laboratory of Cell Biology

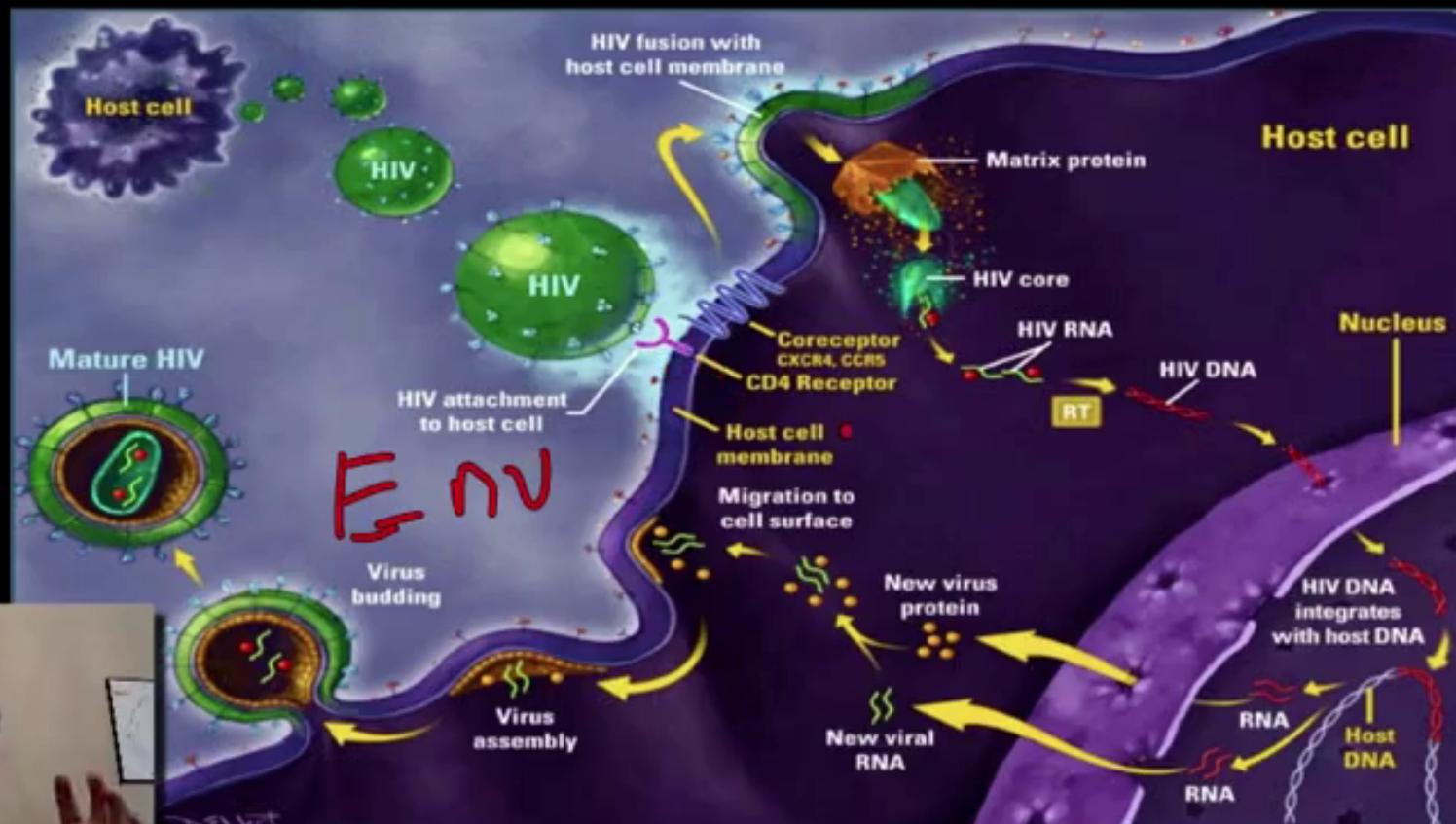
Center for Cancer Research



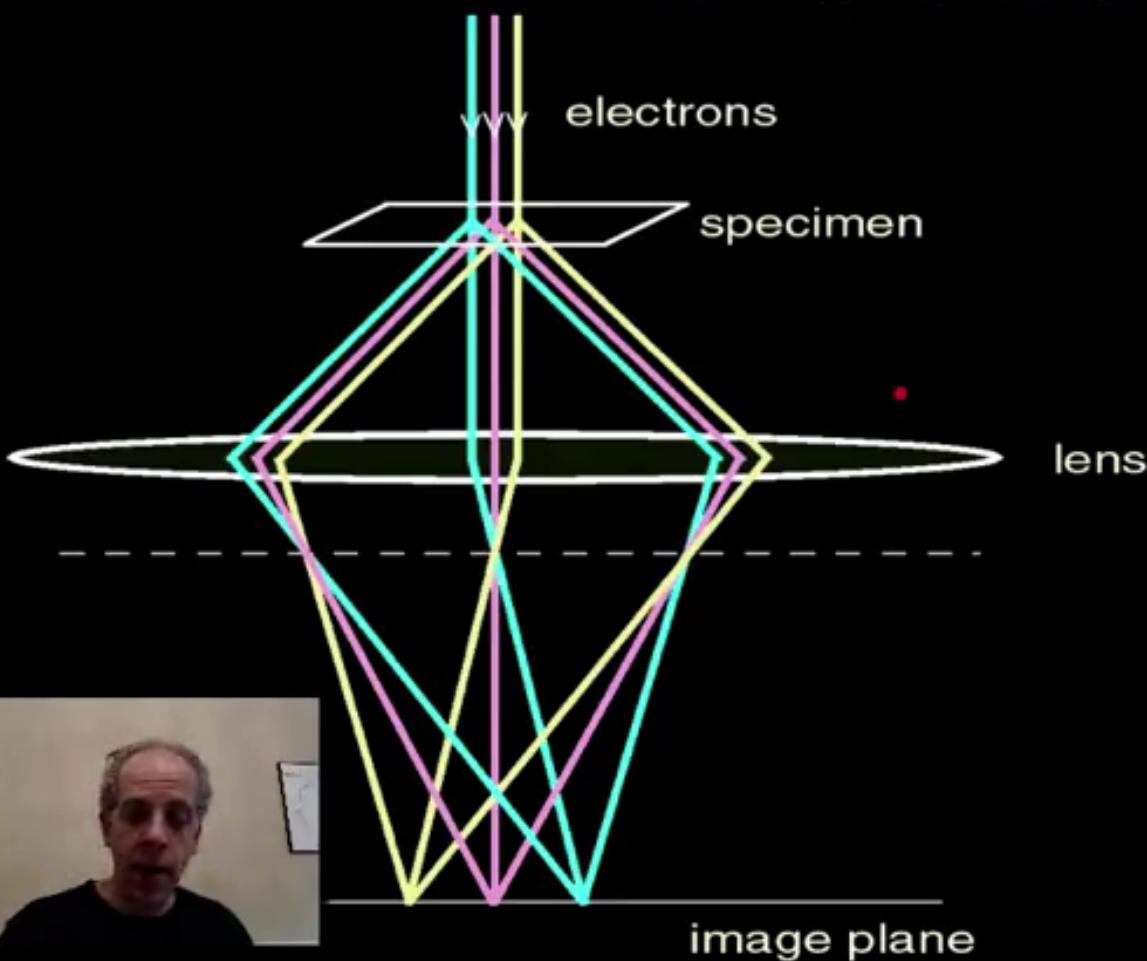
Imaging technologies for biology



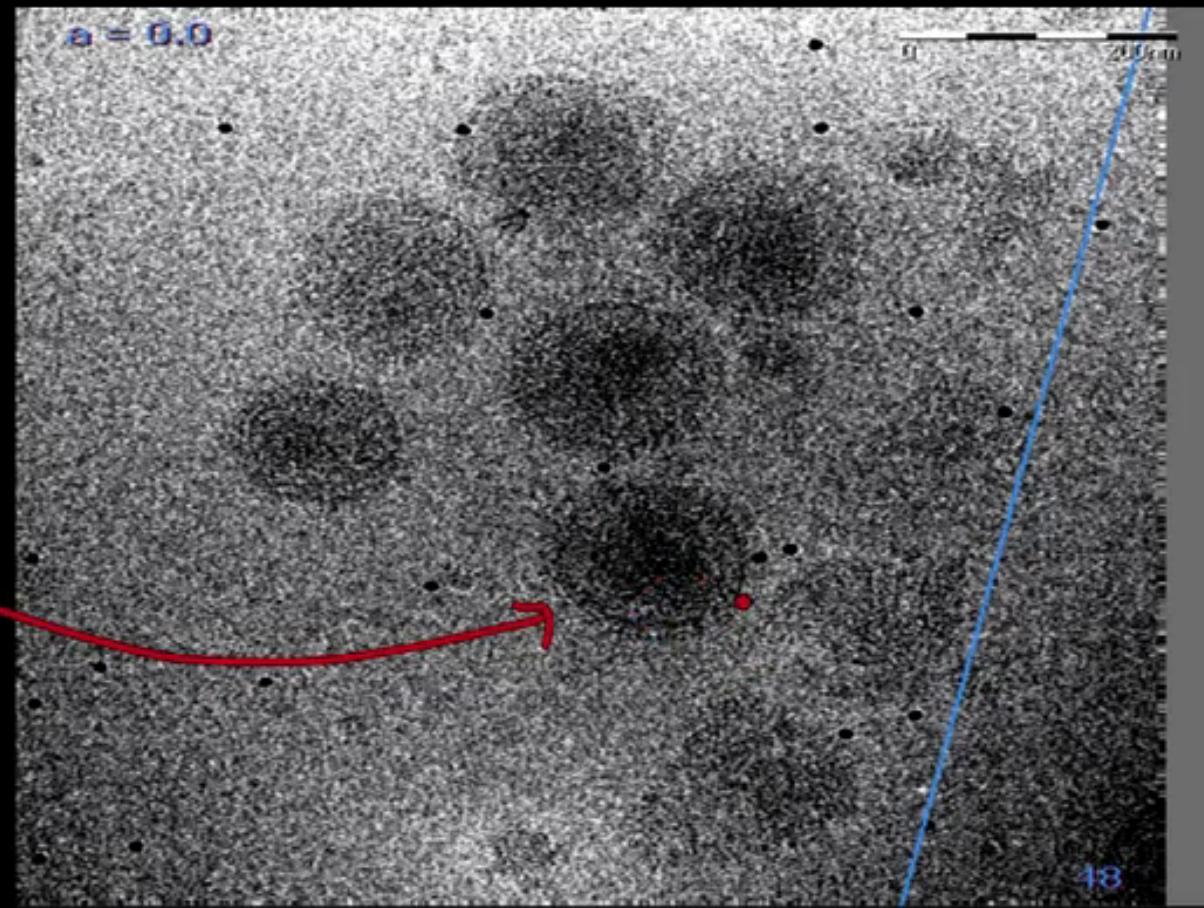
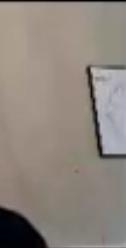
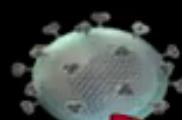
Our Target: Molecular structure of HIV envelope glycoproteins



Transmission Electron Microscopy

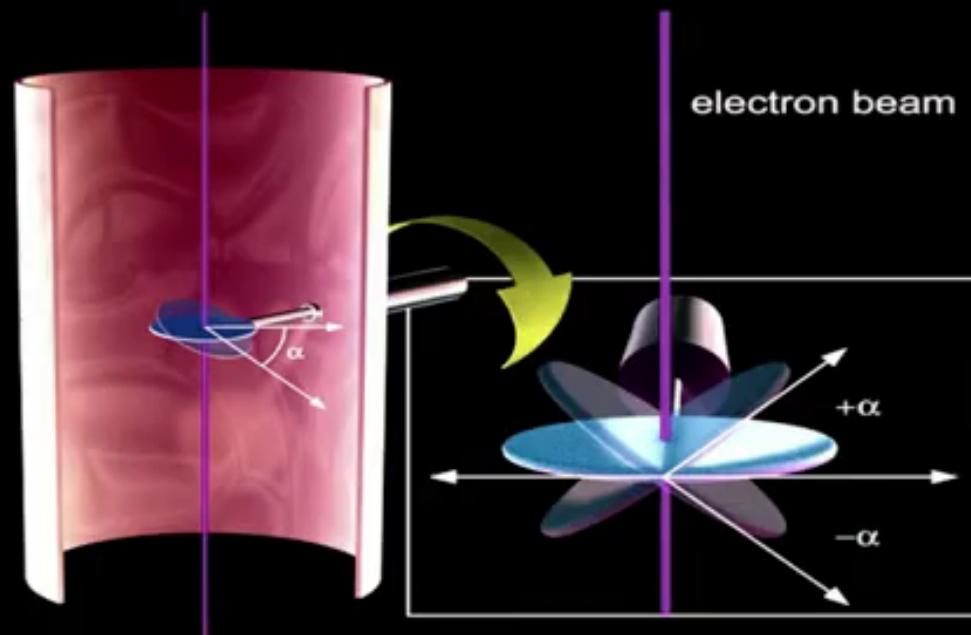


Single Projection Image of HIV

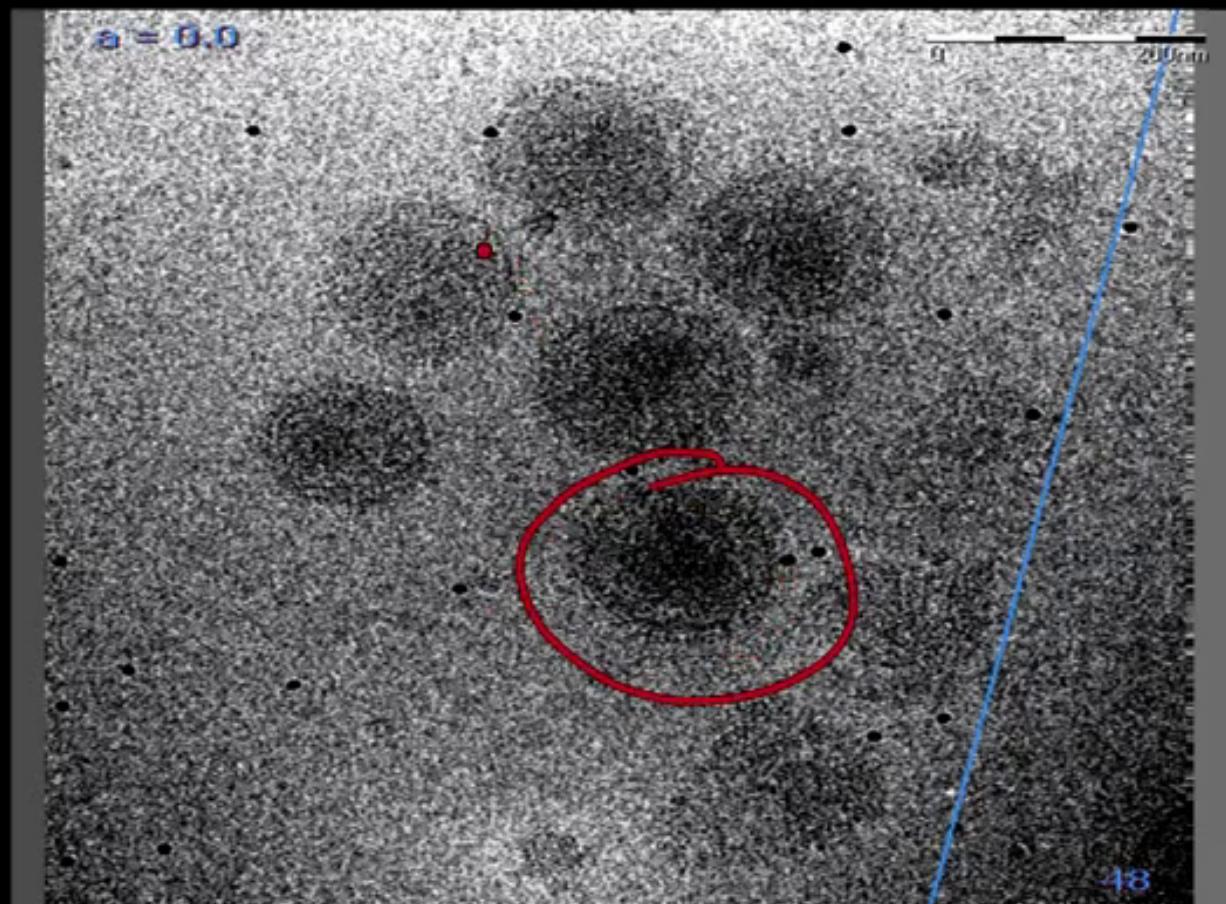


Cryo-Electron Tomography

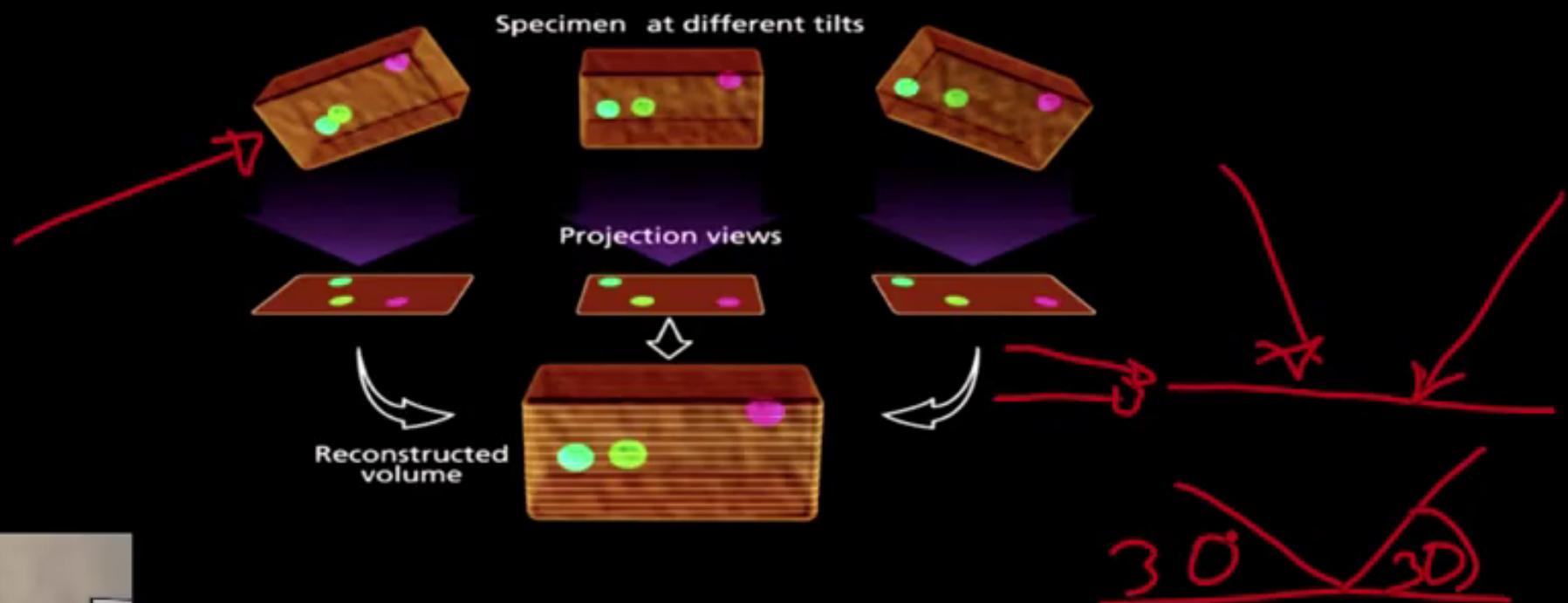
- Reduce radiation damage
- Obtain 3D information



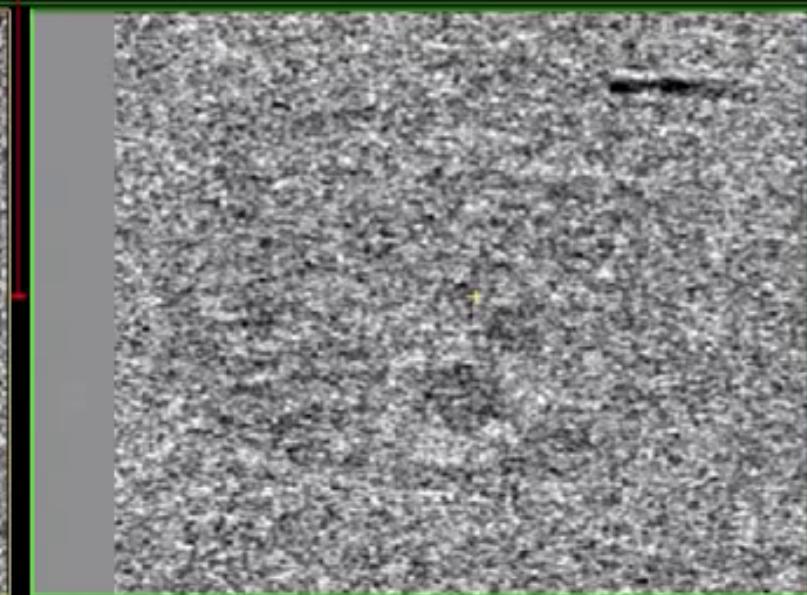
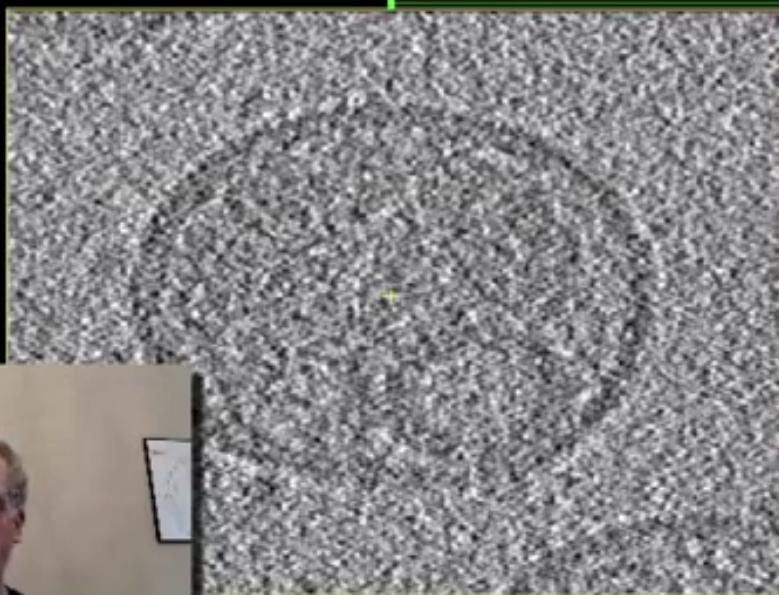
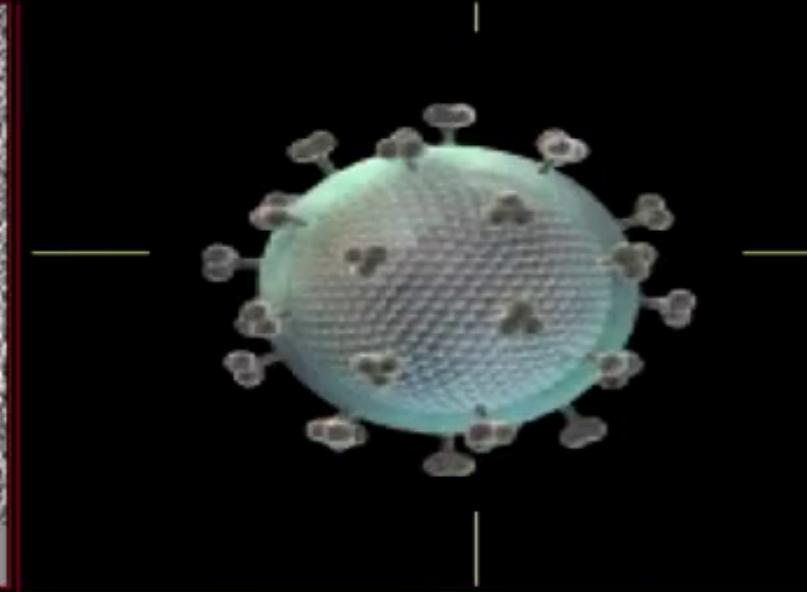
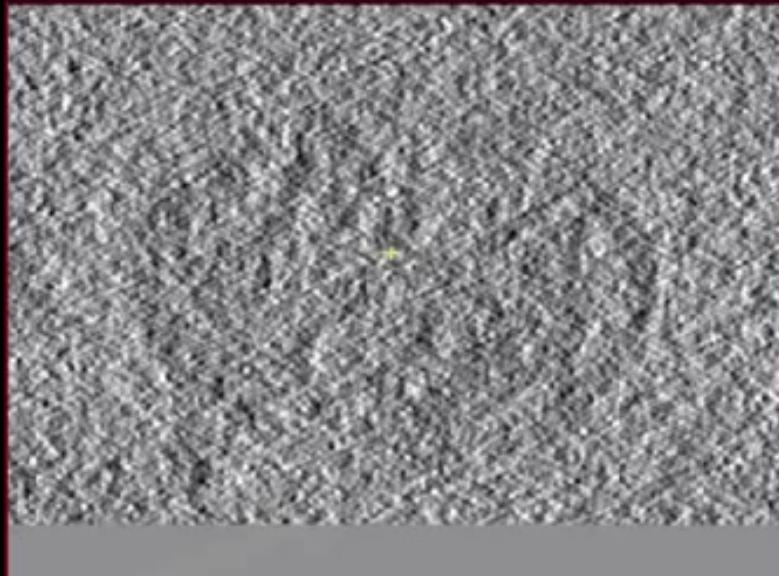
Raw tilt-series of HIV



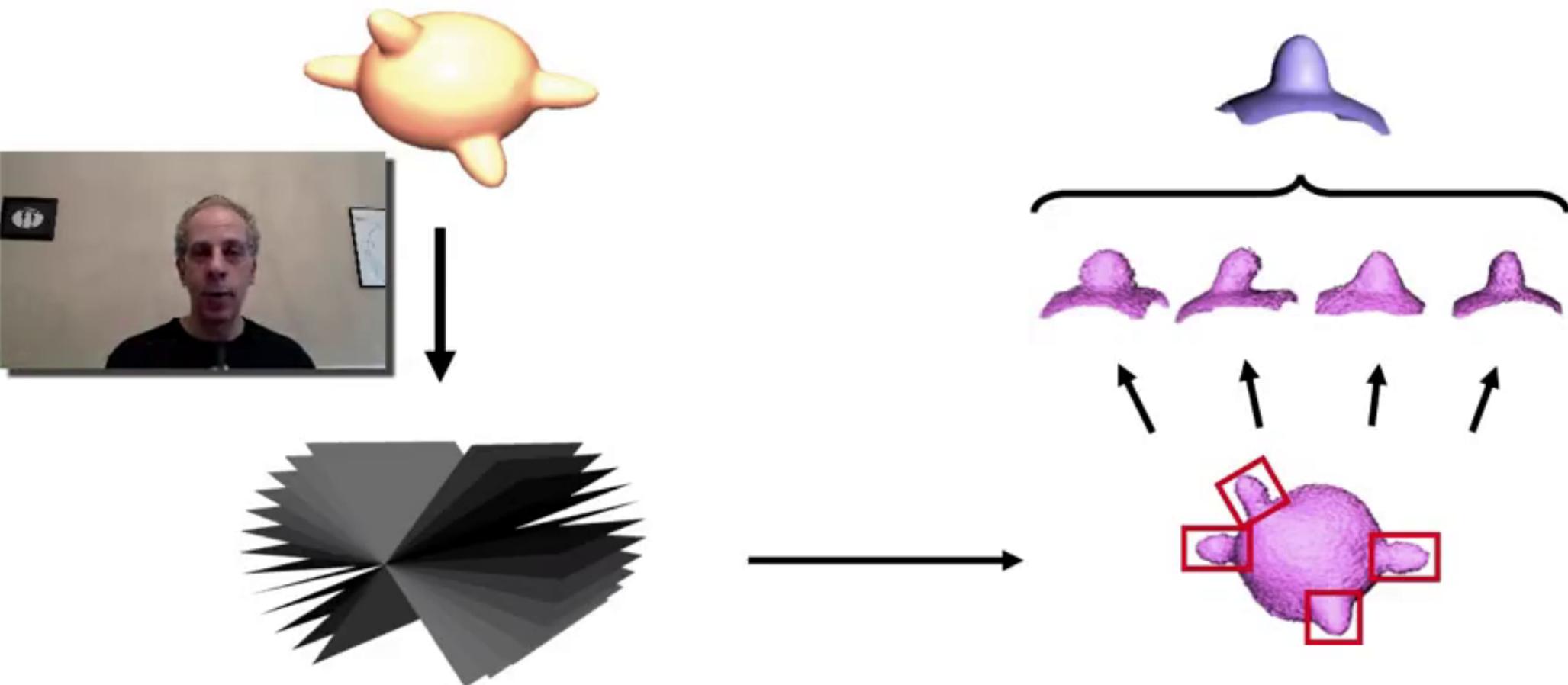
Tomographic Reconstruction



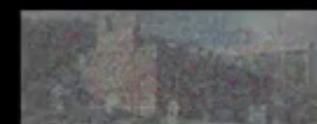
Subramaniam et al., ASM News 60, 240-245.



Sub-Volume Averaging in Electron Tomography



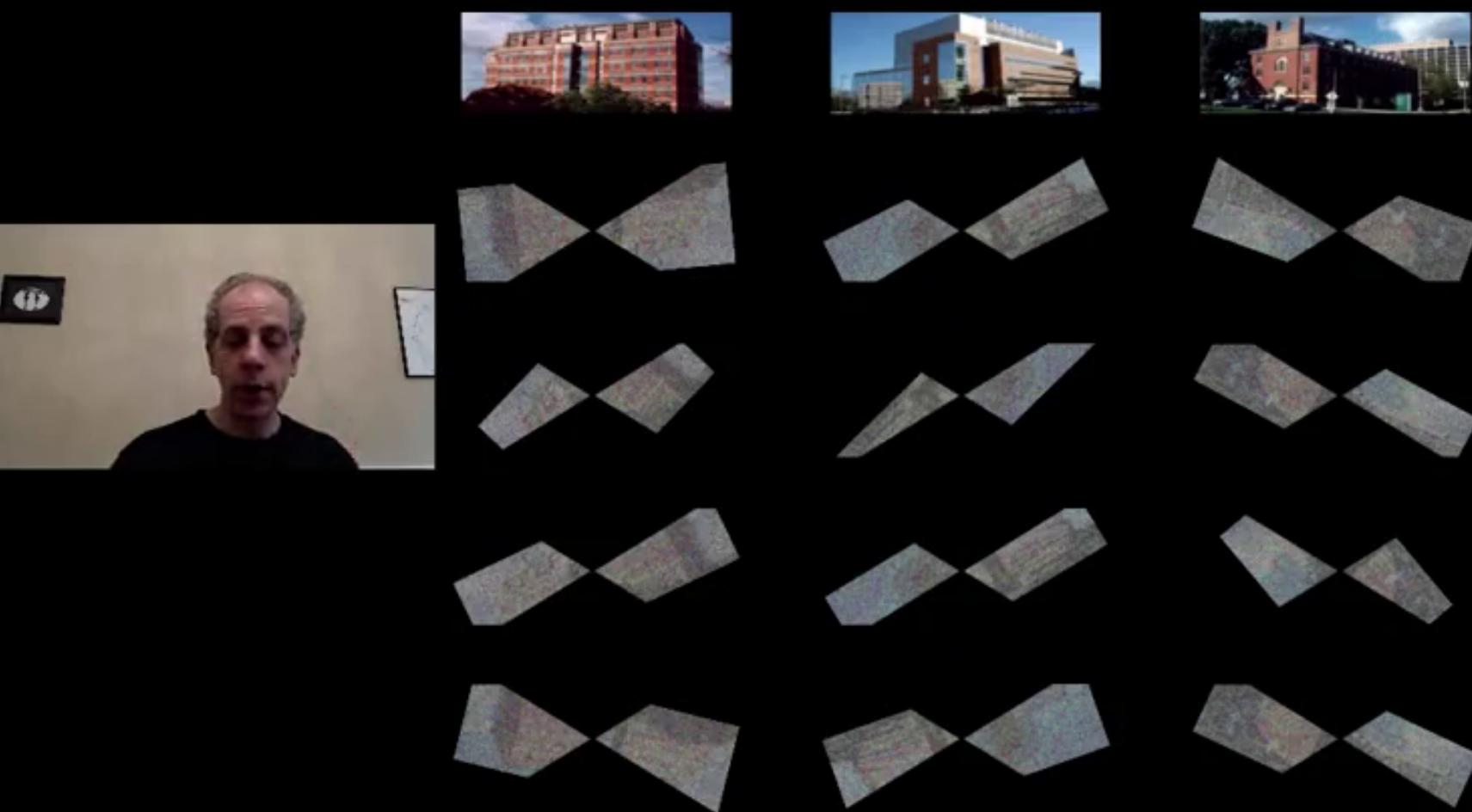
Reconstitution From Noisy and Incomplete Images



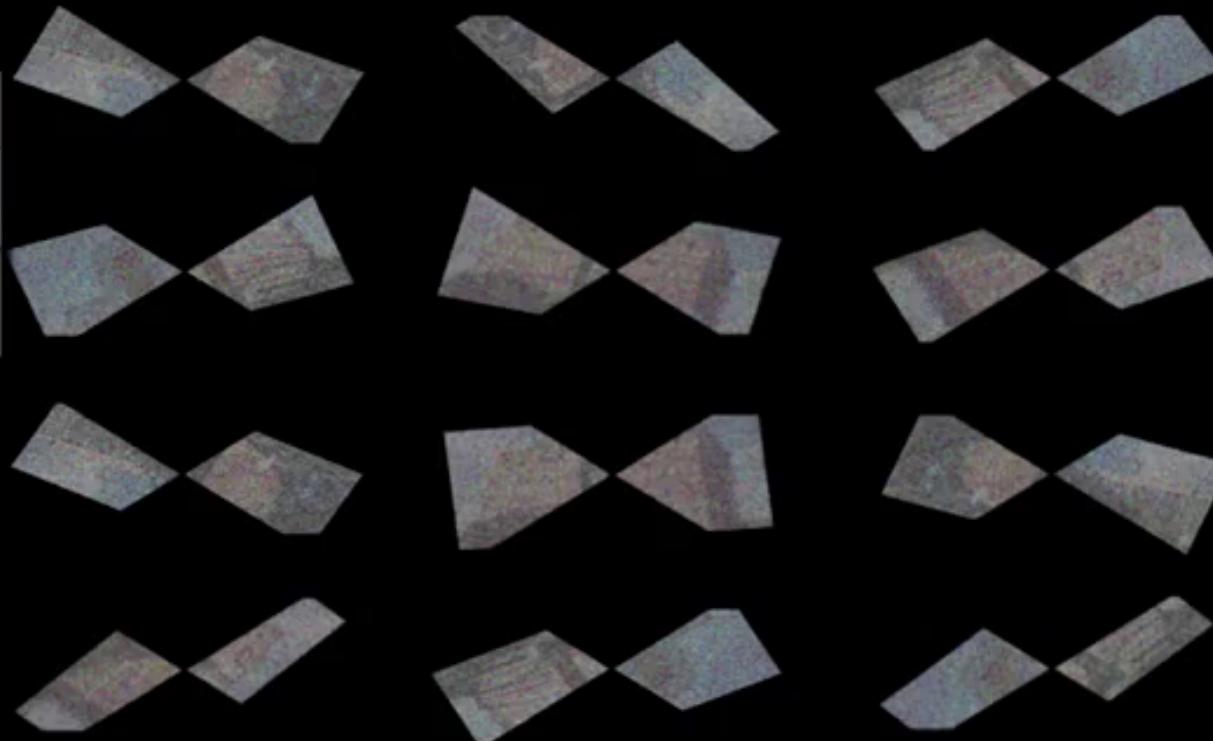
Reconstitution From Noisy and Incomplete Images



Reconstitution From Noisy and Incomplete Images



Reconstitution From Noisy and Incomplete Images



Imaging Challenges of Sub-Volume Averaging in ET

- Low SNR makes alignment difficult
- Alignment ambiguities due to missing data
- 3D datasets require extensive computation

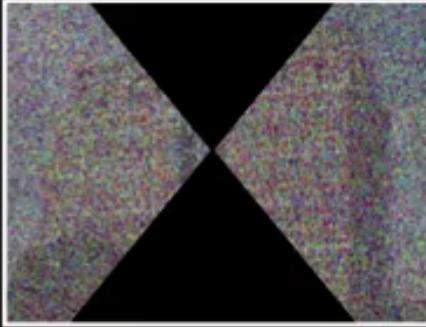
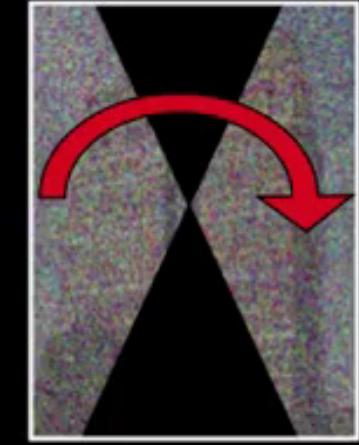


Effects of Missing Data on Alignment

$d(\text{[image]}, \text{[image]})$

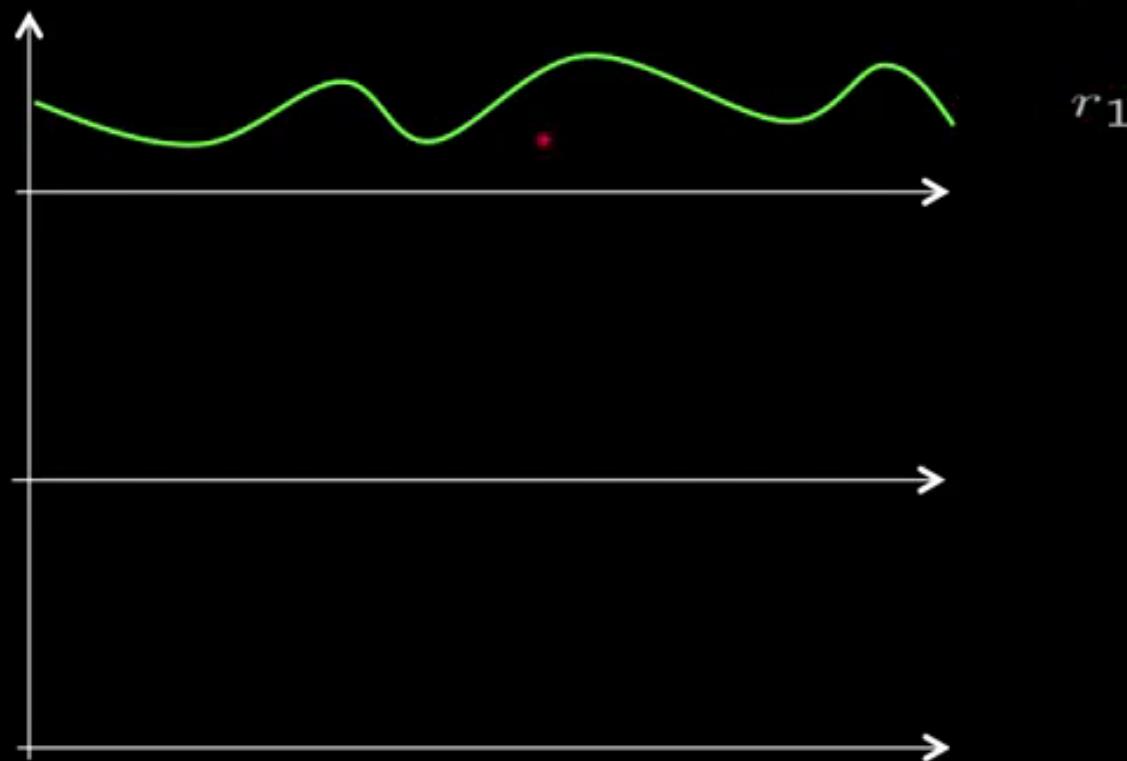


Effects of Missing Data on Alignment

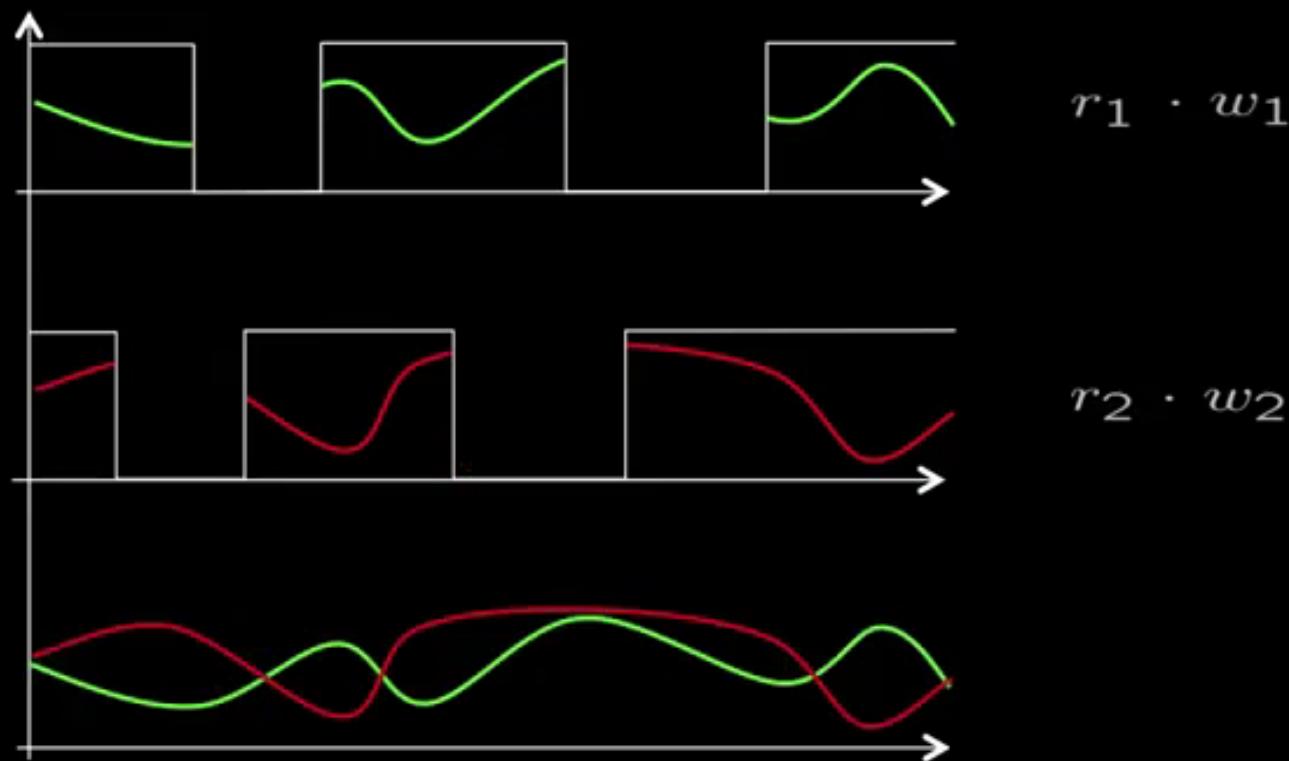
$d($  , )



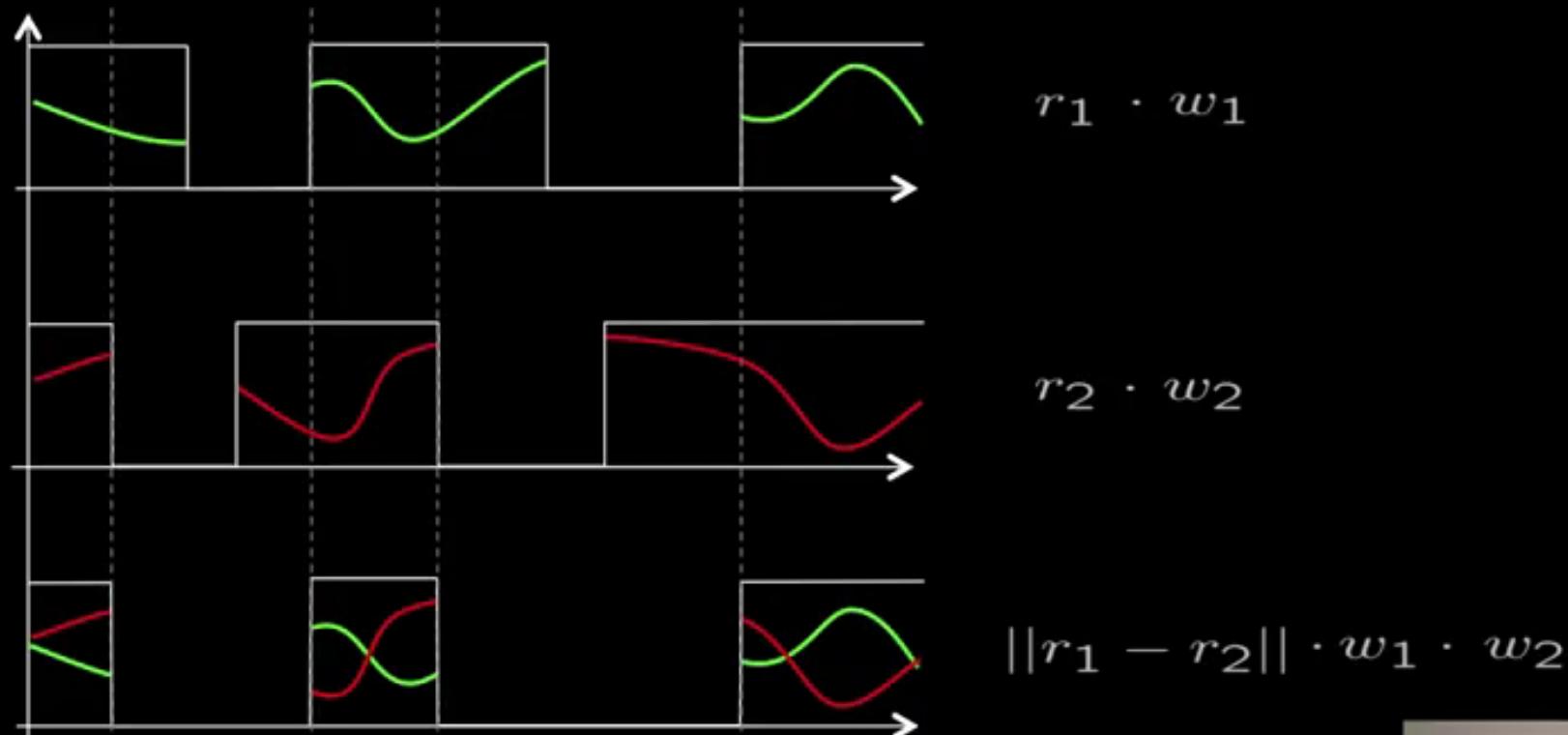
Similarity with Missing Information



Similarity with Missing Information



Similarity with Missing Information

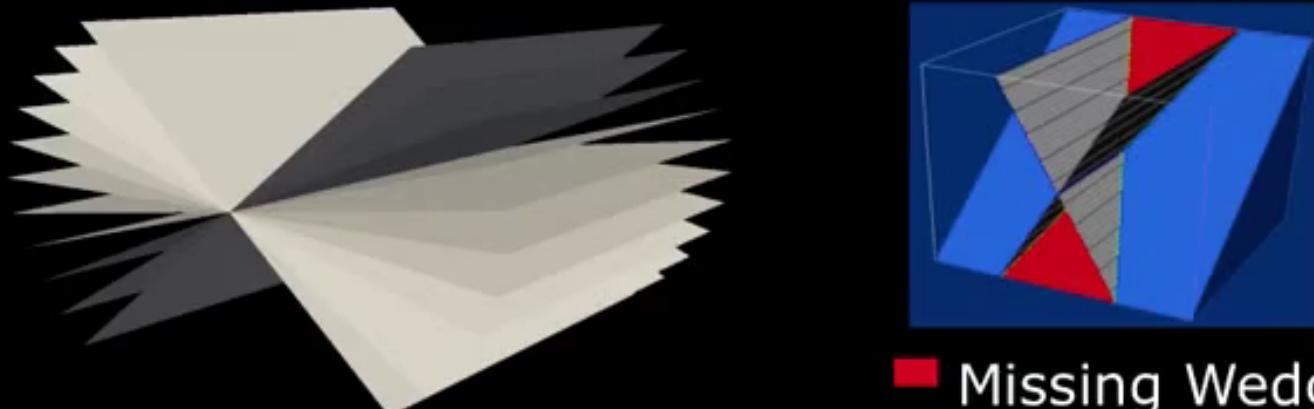


$$d = \frac{\sum \|r_1 - r_2\| \cdot w_1 \cdot w_2}{\sum w_1 \cdot w_2}$$



Similarity of partially occluded volumes in Fourier space

$$\hat{\mathcal{F}}_1 = \mathcal{F}_1 \mathcal{W}_1, \quad \hat{\mathcal{F}}_2 = \mathcal{F}_2 \mathcal{W}_2, \quad \mathcal{W}_i \rightarrow [0, 1]$$

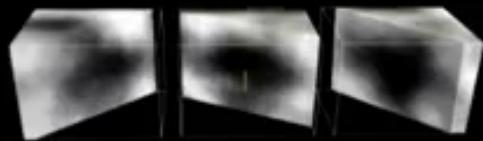


$$d = \frac{\int_{\mathcal{B}} ||\hat{\mathcal{F}}_1 - \hat{\mathcal{F}}_2|| \mathcal{W}_1 \mathcal{W}_2}{\int_{\mathcal{B}} \mathcal{W}_1 \mathcal{W}_2}$$



1000

image Optimization Strategy

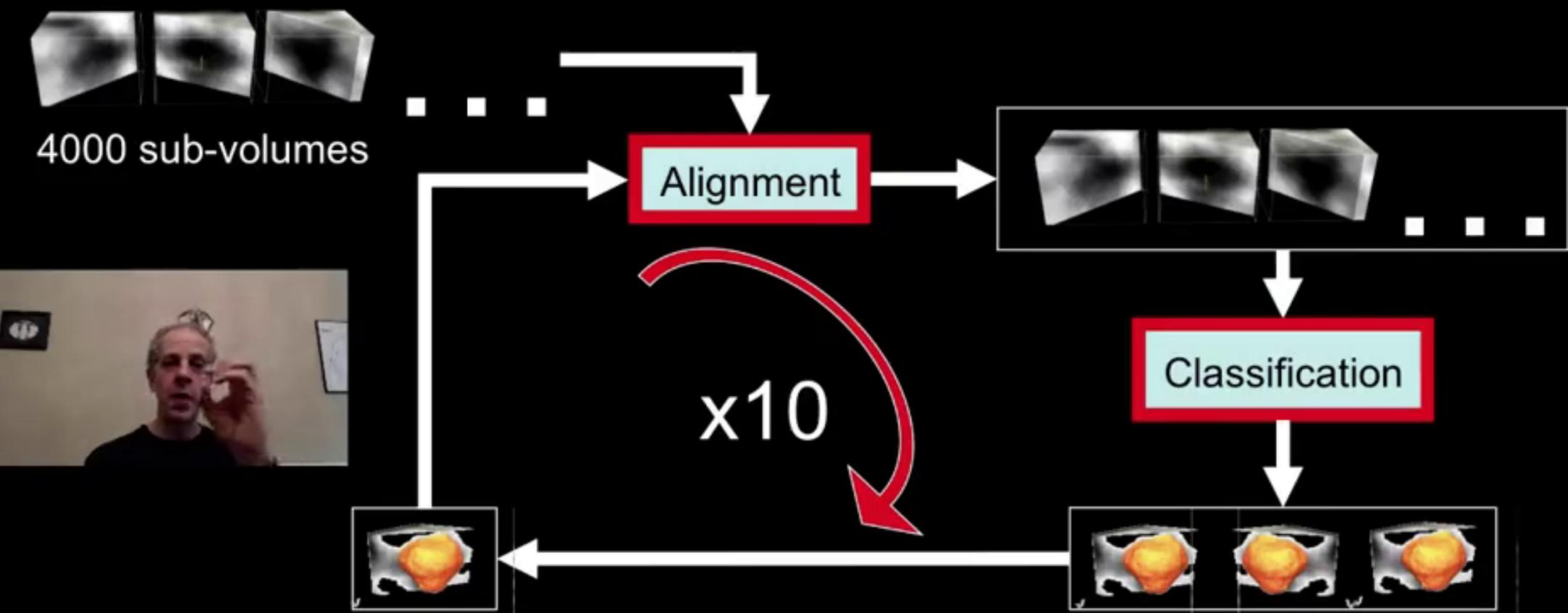


4000 sub-volumes

...



Image Optimization Strategy



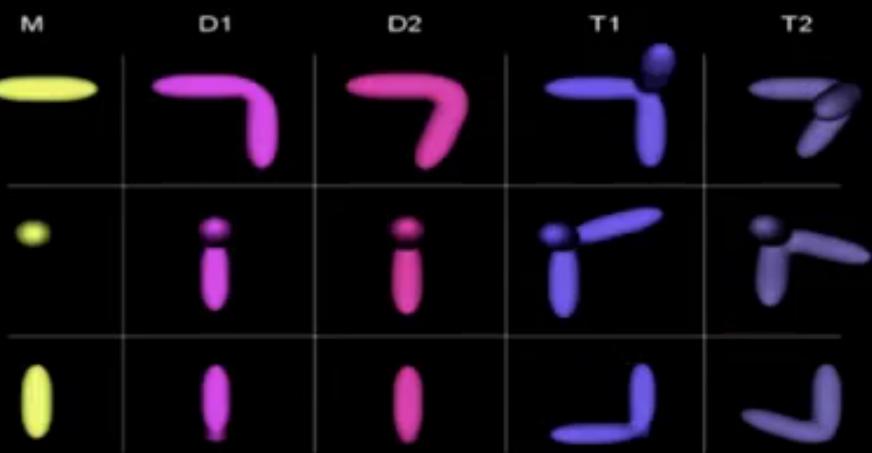
3D Image Alignment



$$\min_{\theta, \phi, \psi, \vec{t}} d(\text{3D Model}, \text{Image})$$



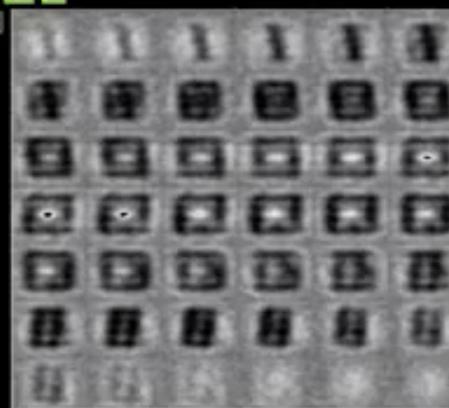
- 6 DOF problem: Speed-up in Fourier domain



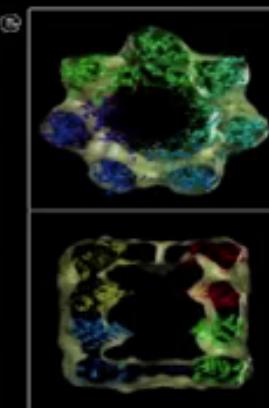
Phantoms



GroEL



d



e



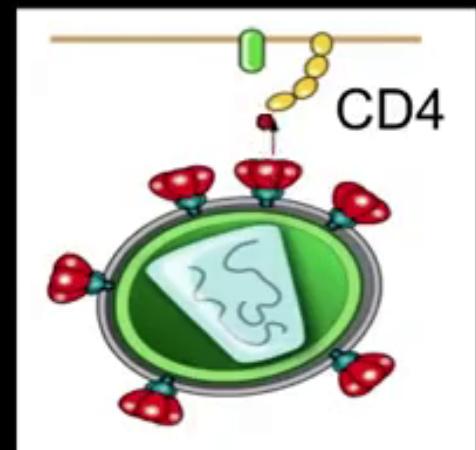
HIV





HIV envelope glycoproteins

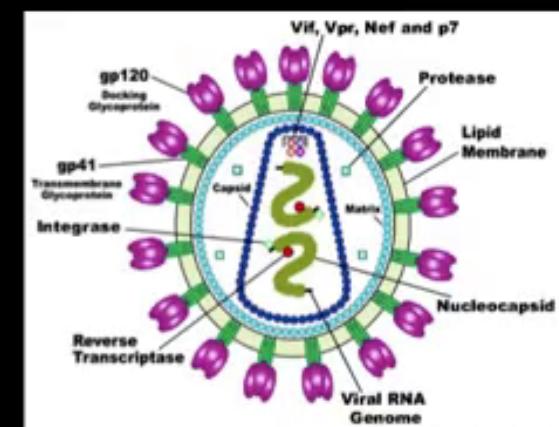
- Mediate virus binding to the cell surface receptor CD4 on target cells to initiate infection
- Functional unit is a trimer



of **gp120** (surface glycoprotein)

and **gp41** (transmembrane unit).

- Structure of components available.
- Structure of the trimer remains elusive.





38/nature07159

nature

LETTERS

Molecular architecture of native HIV-1 gp120 trimers

Jun Liu¹*, Alberto Bartesaghi¹*, Mario J. Borgnia¹*, Guillermo Sapiro² & Sriram Subramaniam¹

¹Laboratory of Cell Biology, Center for Cancer Research, National Cancer Institute, NIH, Bethesda, Maryland 20892, USA. ²Department of Electrical and Computer Engineering, University of Minnesota, Minneapolis, Minnesota 55455, USA.

*These authors contributed equally to this work.

- Use cryo-electron tomography combined with 3D image averaging and classification
- Report 3D “snapshots” of trimeric spike:
 - Unliganded state
 - Complex with broadly neutralizing b12
 - Ternary complex with CD4 and 17b

Imaging the spike at different states

- 80 tilt series, 400 virus, 4K spikes
 - 1. Unliganded state
 - 2. Complex with b12
 - 3. Ternary complex with CD4 and 17b



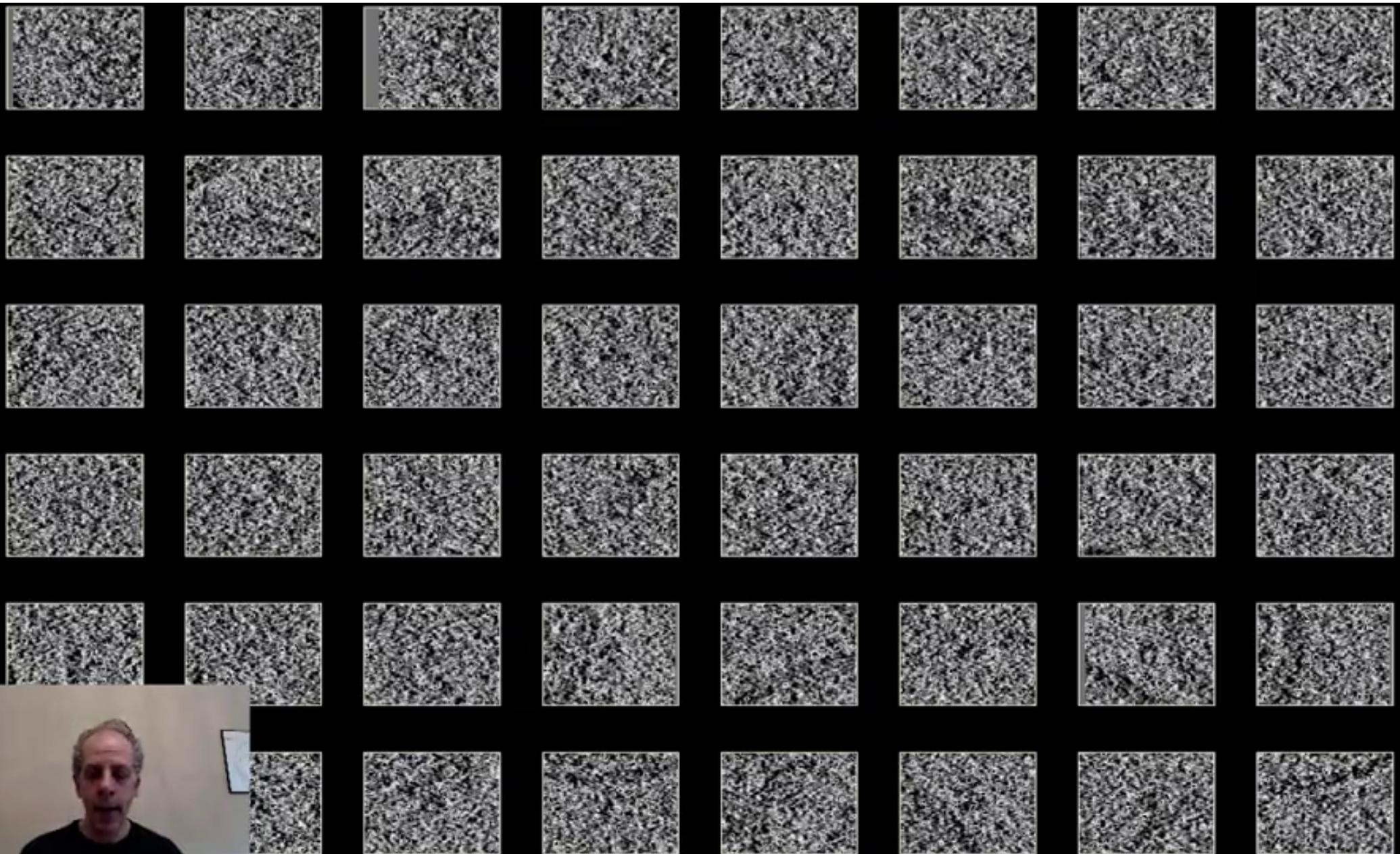


Image Refinement Loop

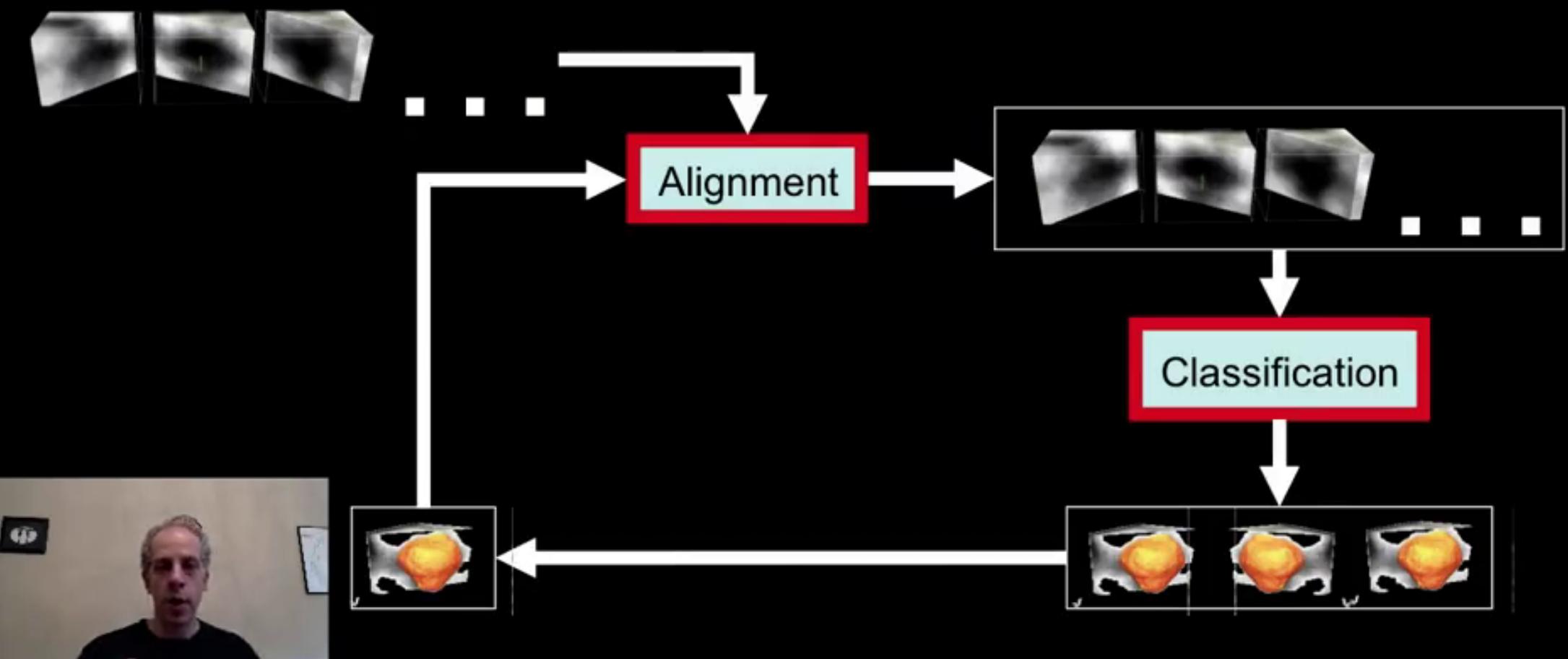
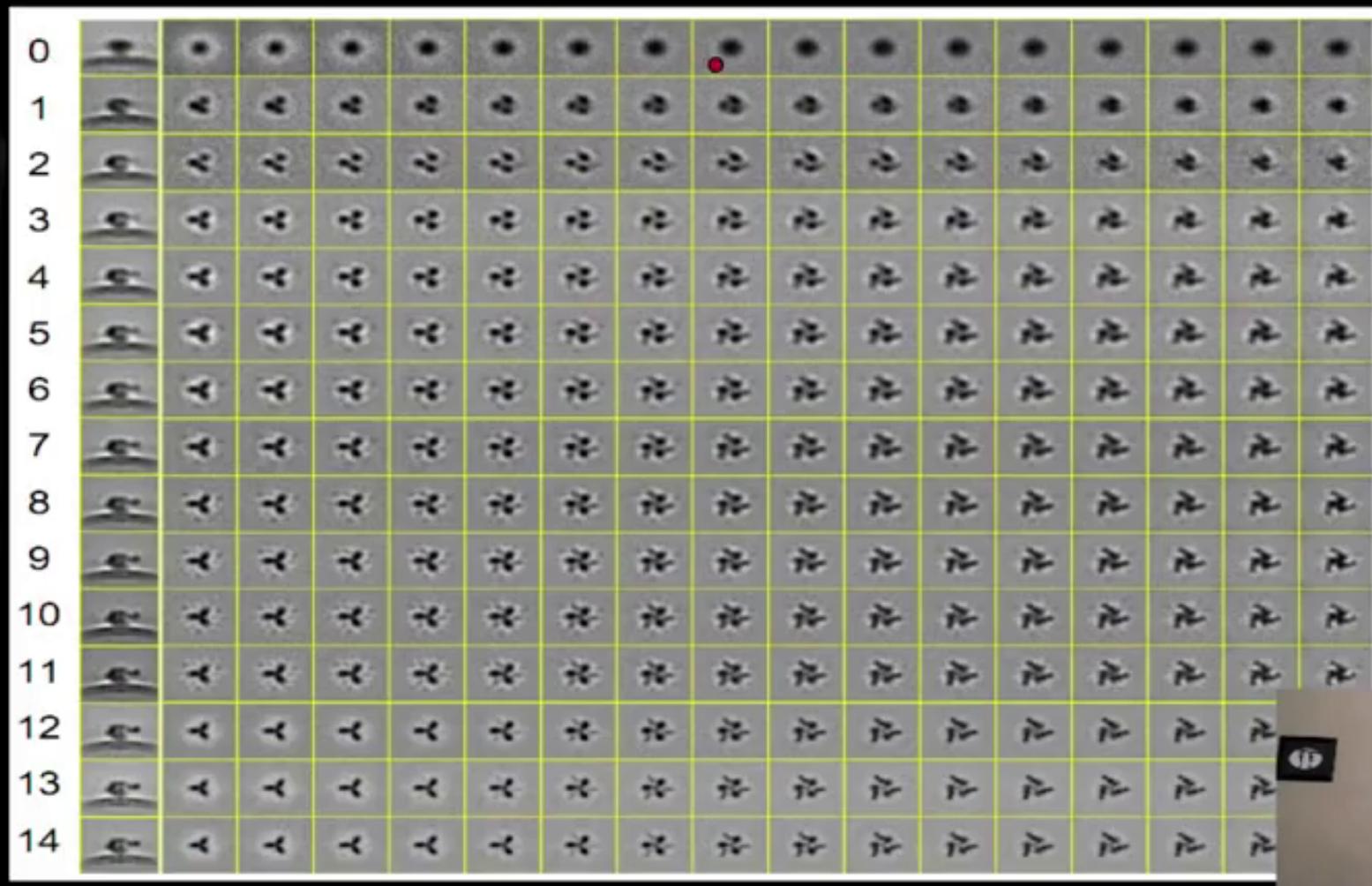
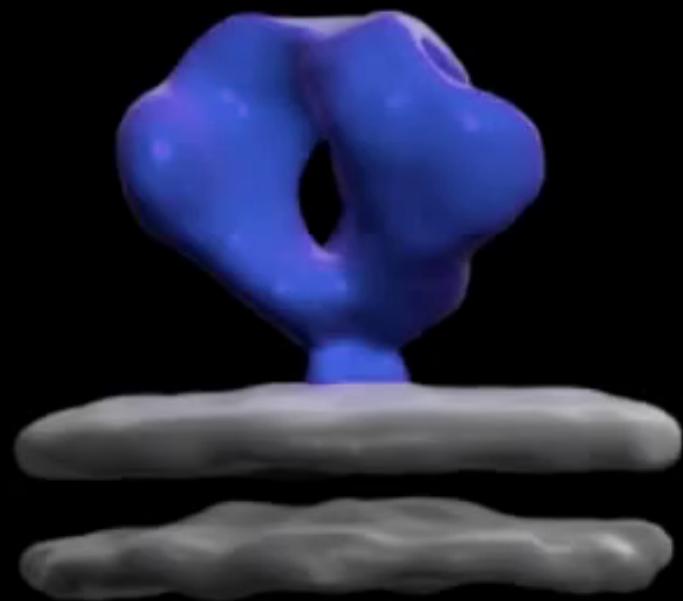
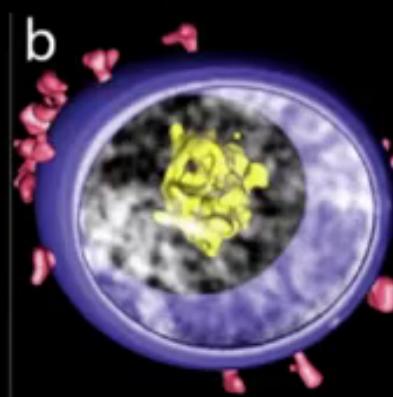
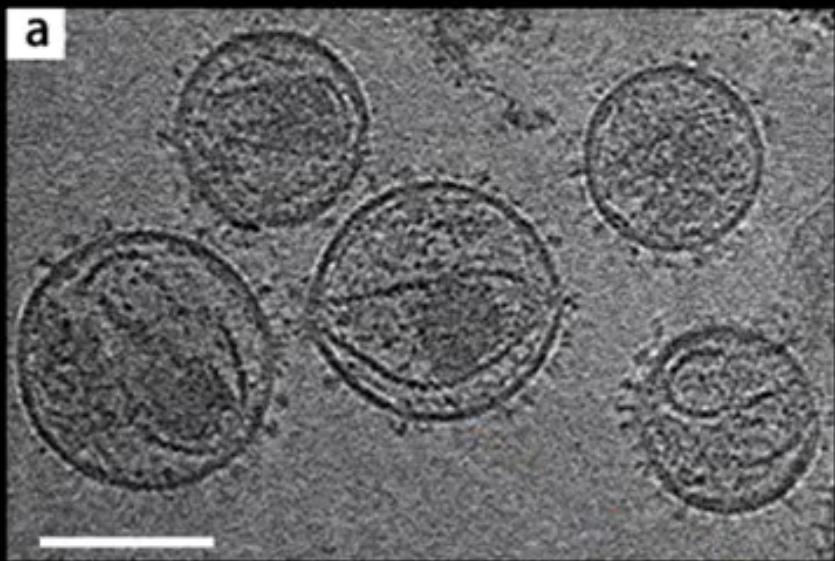
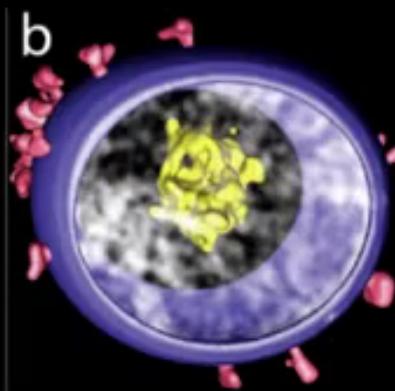
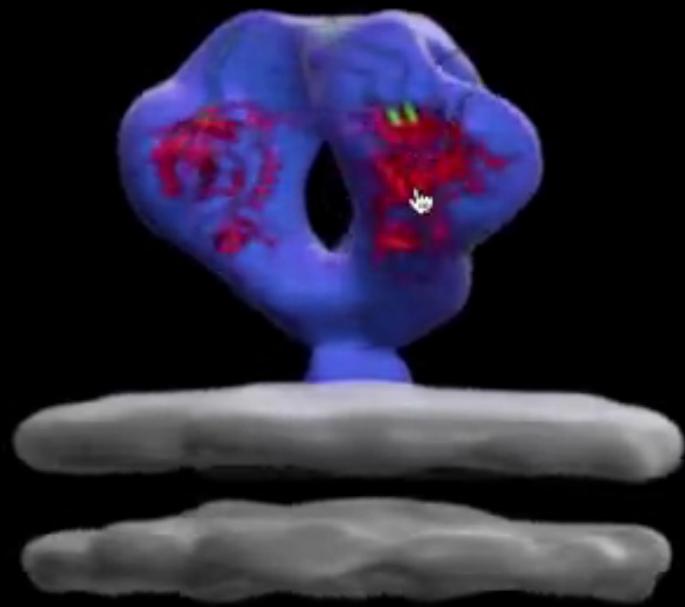
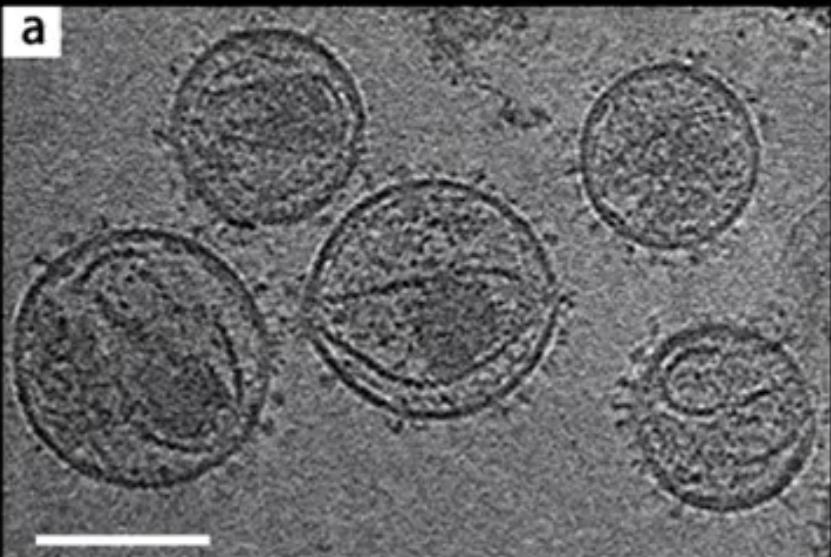


Image Refinement Loop

Refinement Iterations

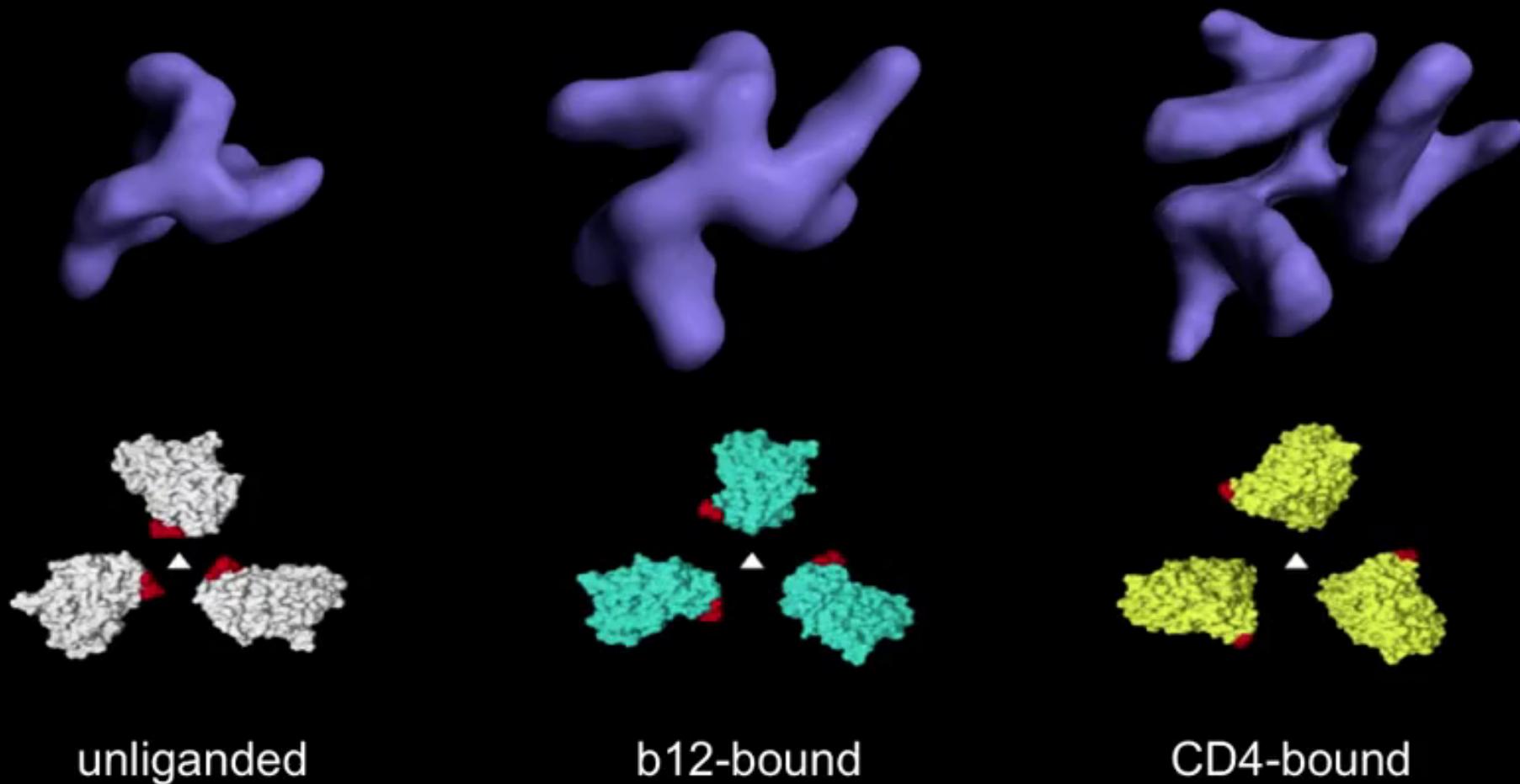




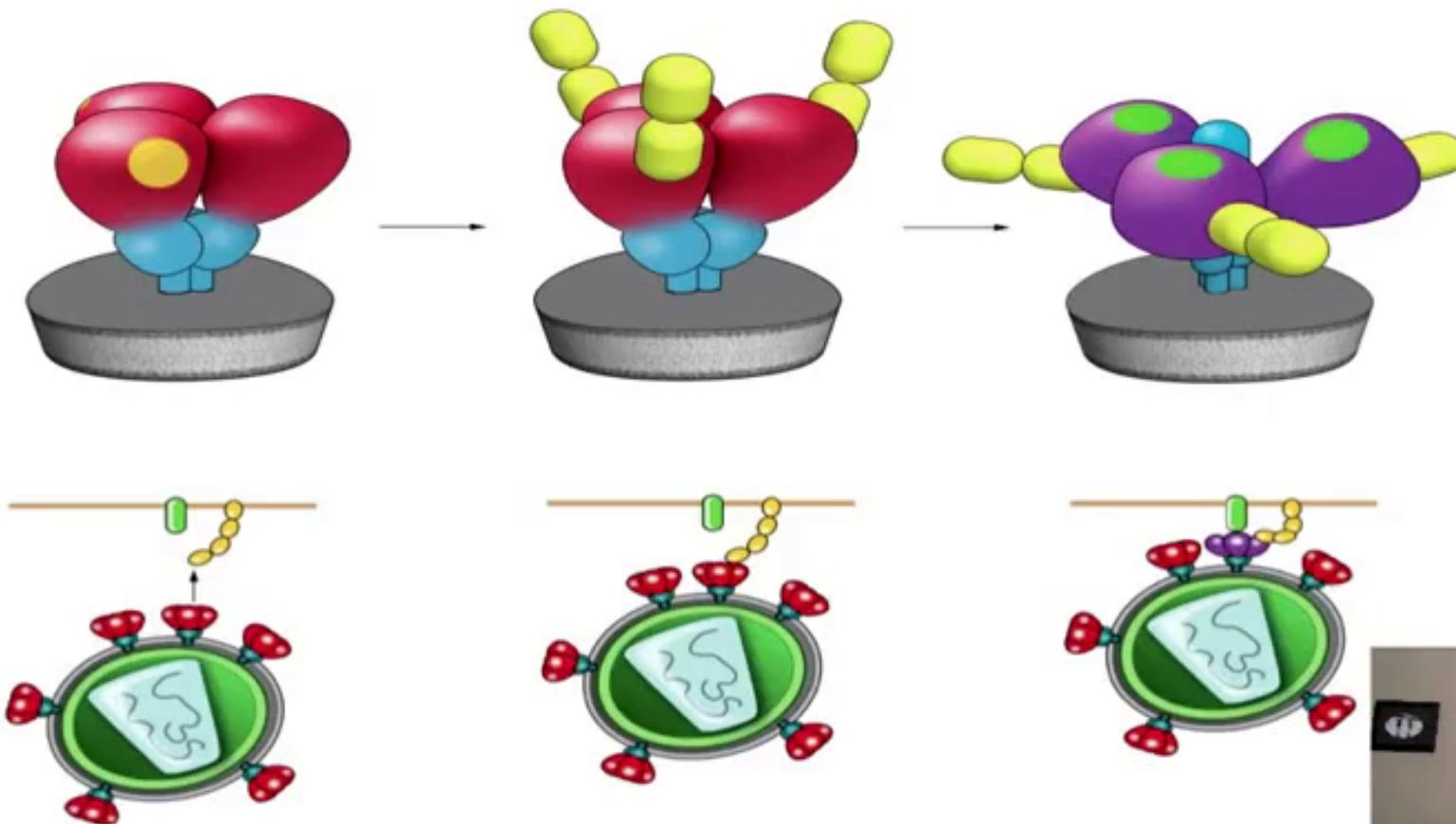




Piecing it all together



Conformational changes of the trimeric spike that occur upon CD4 binding



Diffusion-Weighted MRI

Provides architecture of biological tissues

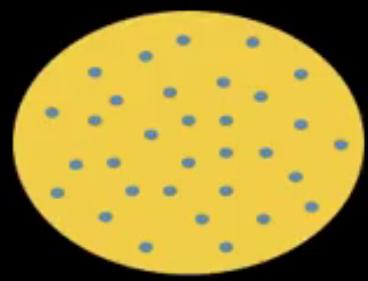


Used to study:

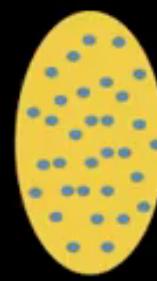
- **Neurological disorders**
- **Brain development**
- **Structure of brain fiber bundles**



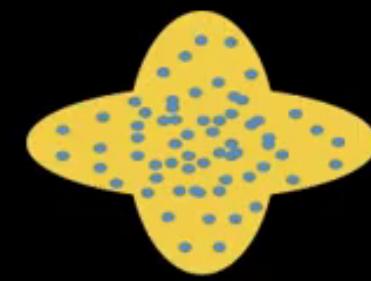
Diffusion of Water Molecules



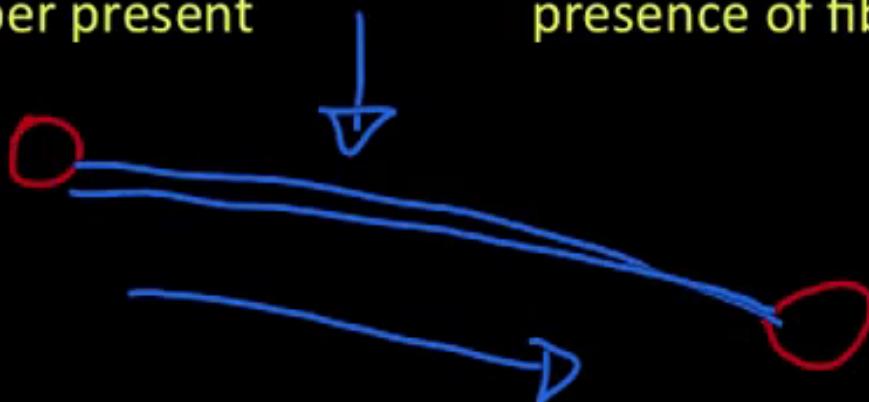
Isotropic Diffusion
no fiber present



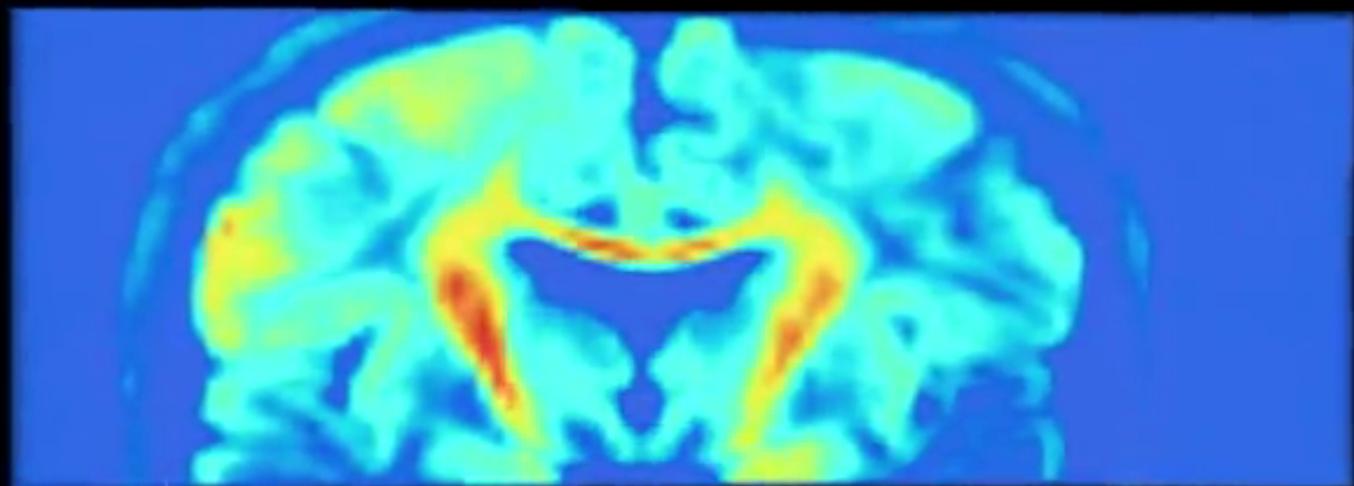
Anisotropic Diffusion
presence of fiber



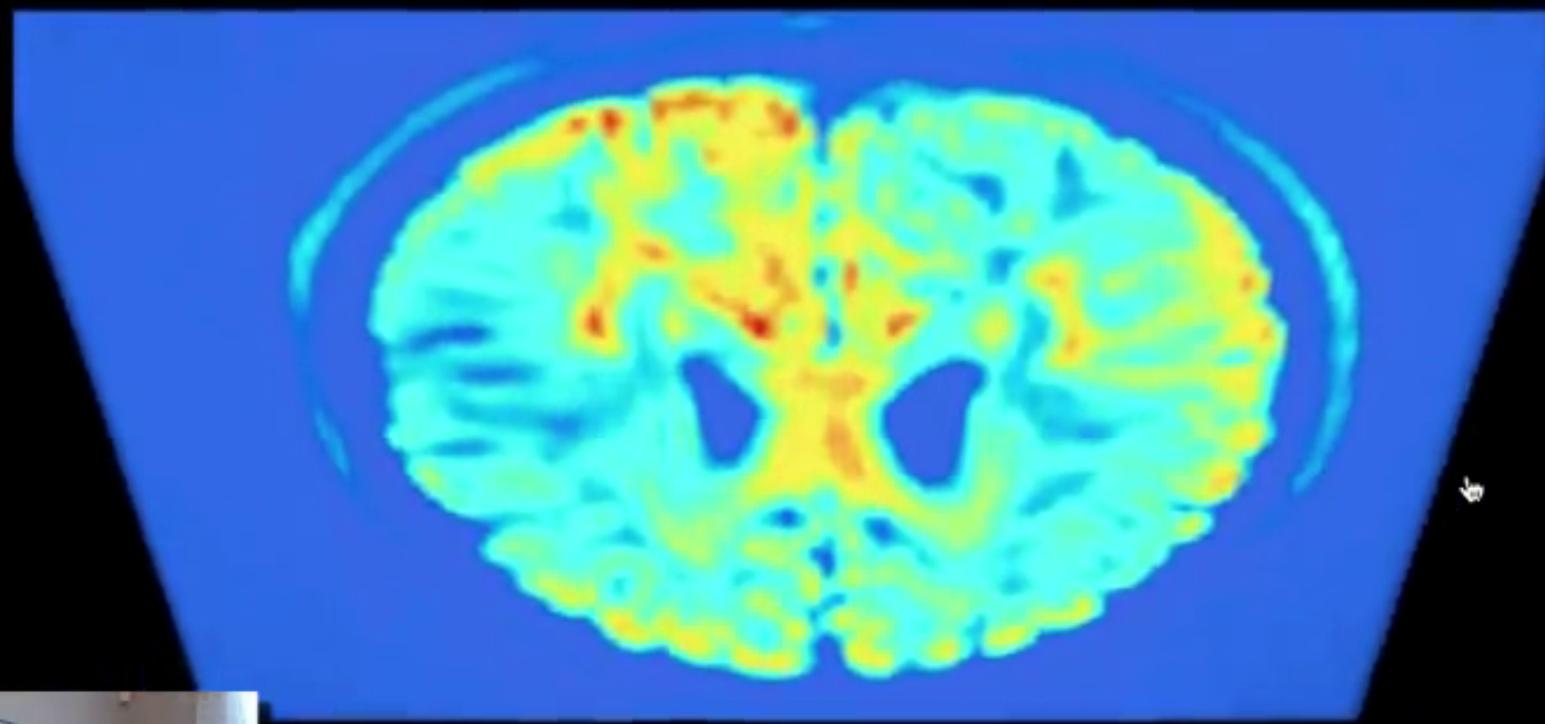
Orthogonal Diffusion
fiber crossing

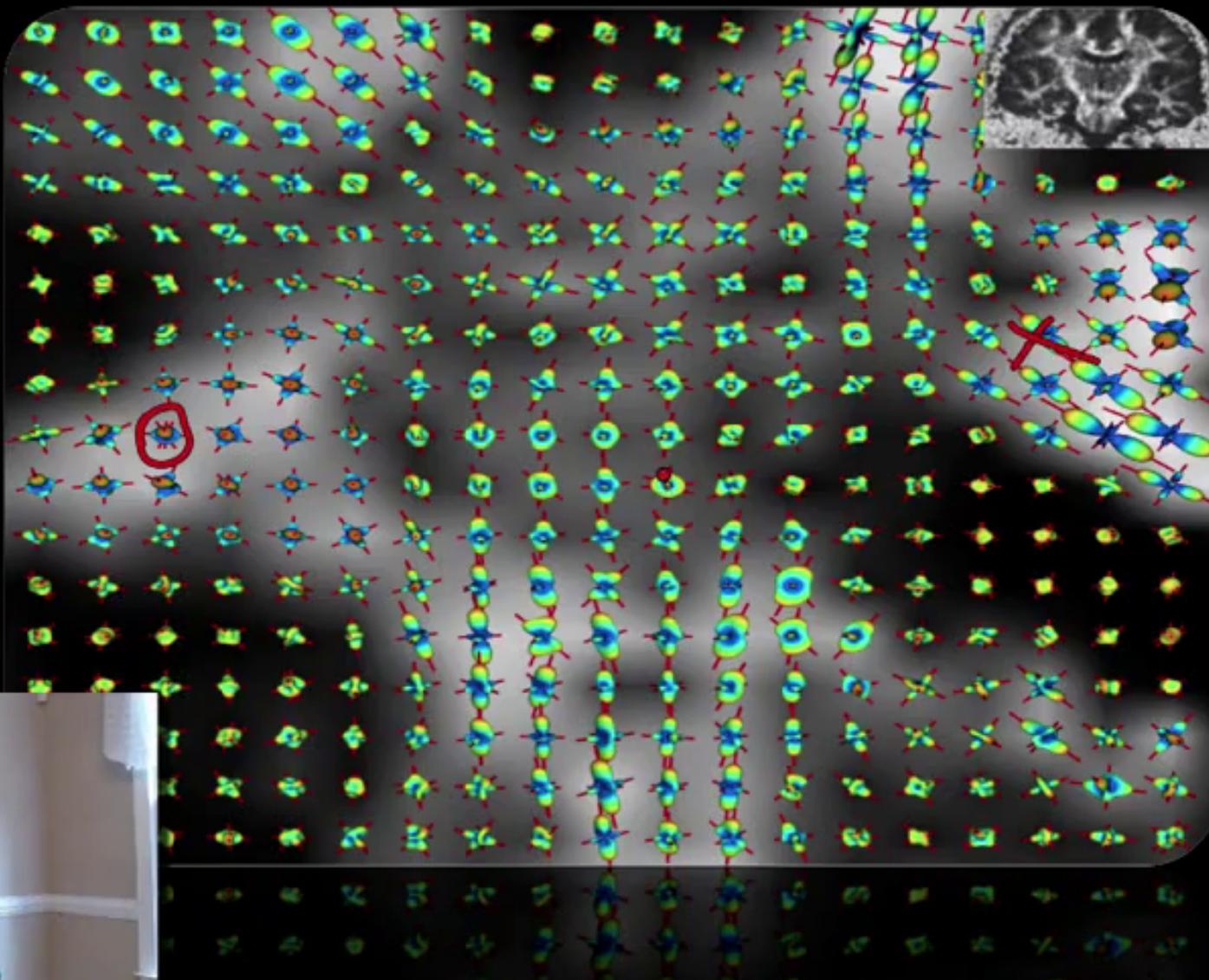


Diffusion-Weighted MR Image

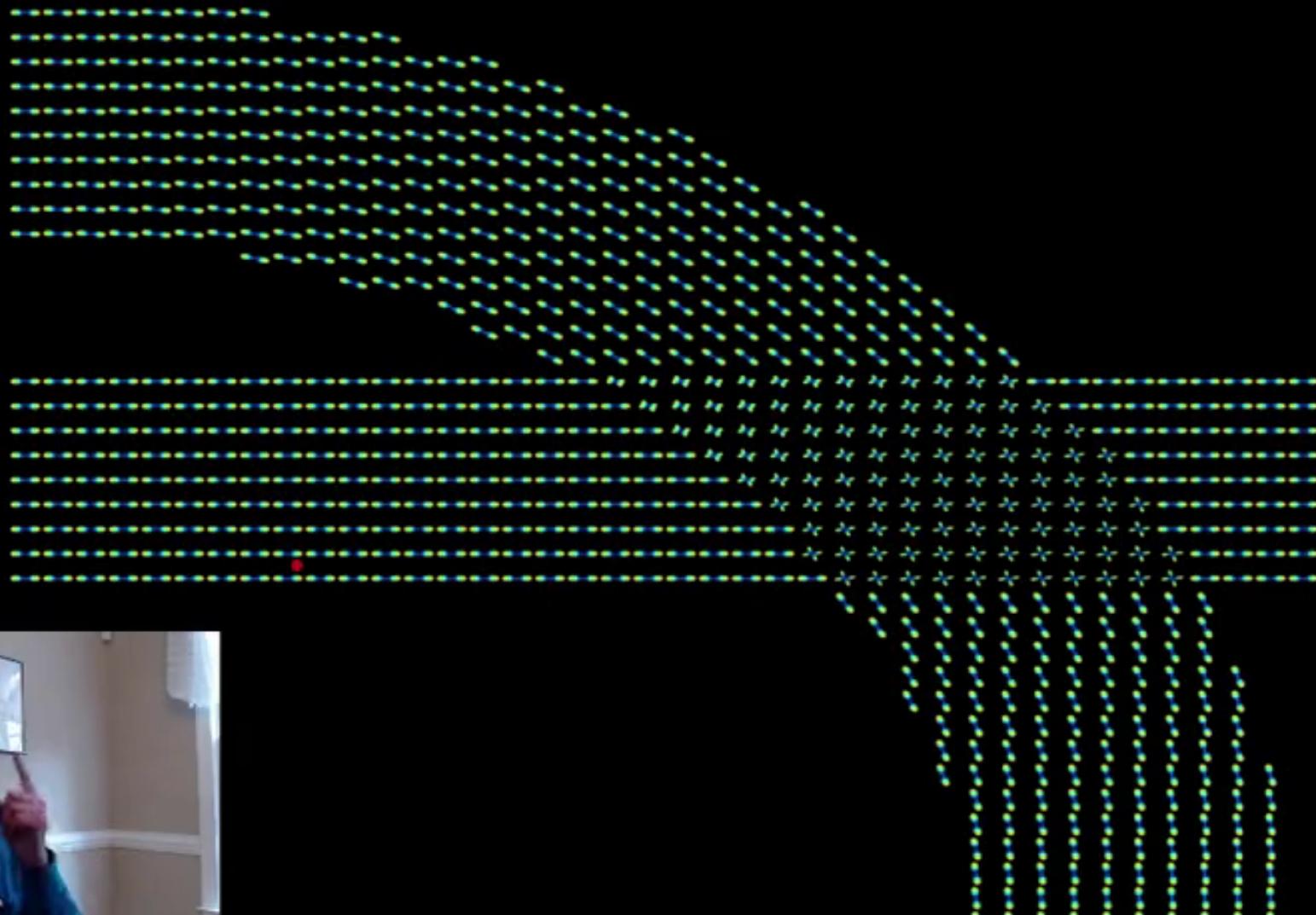


Diffusion-Weighted MR Image

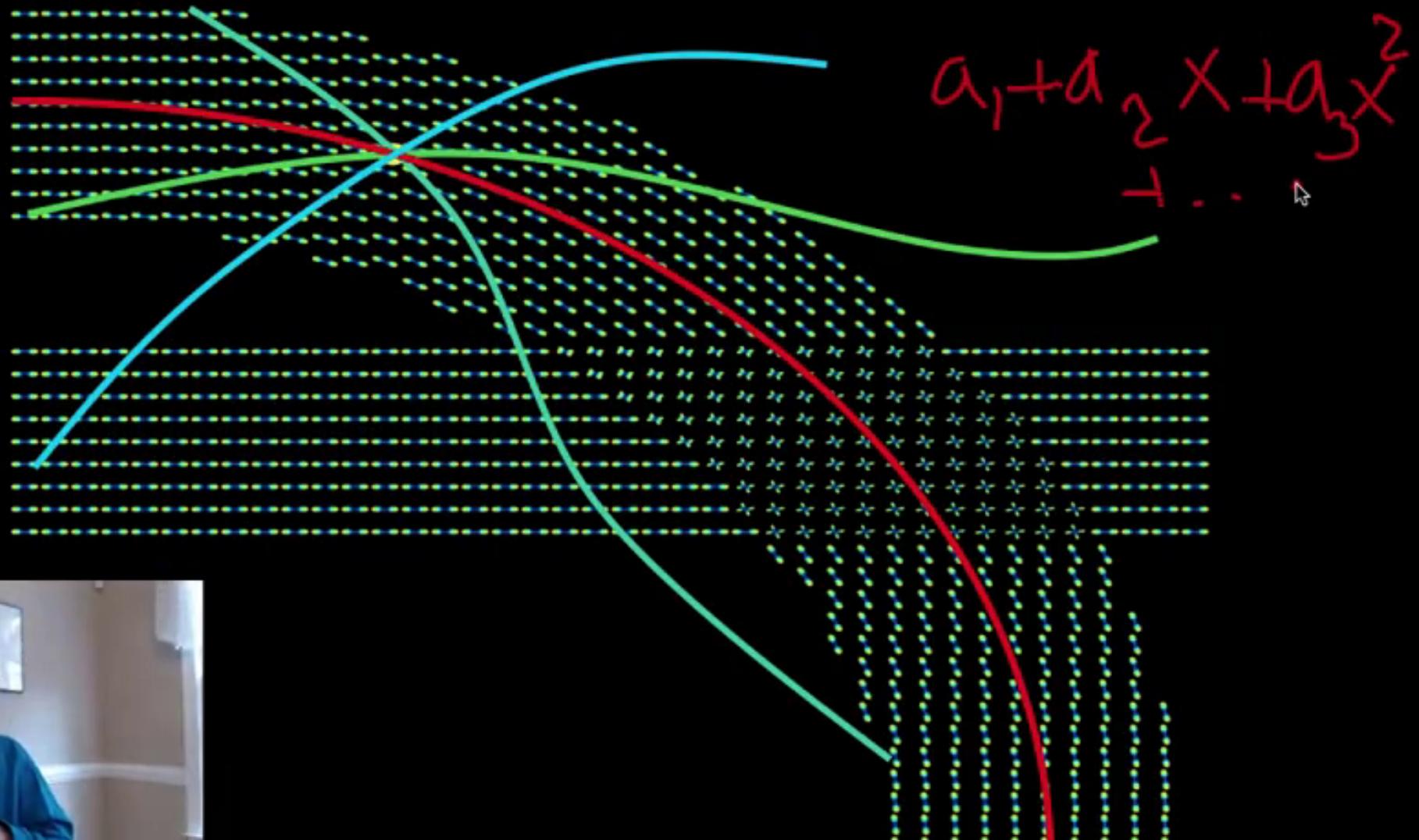




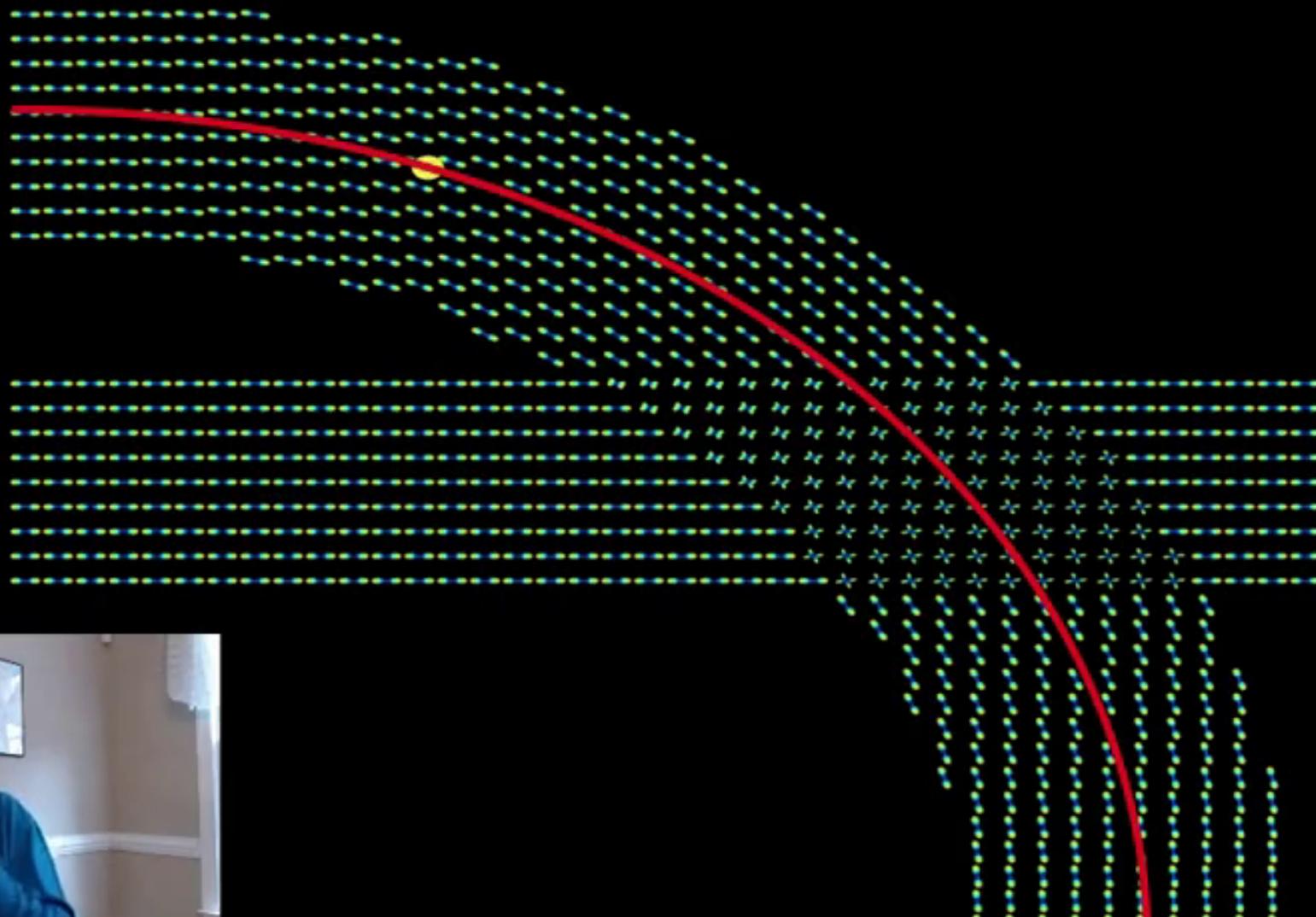
Hough Transform !!!



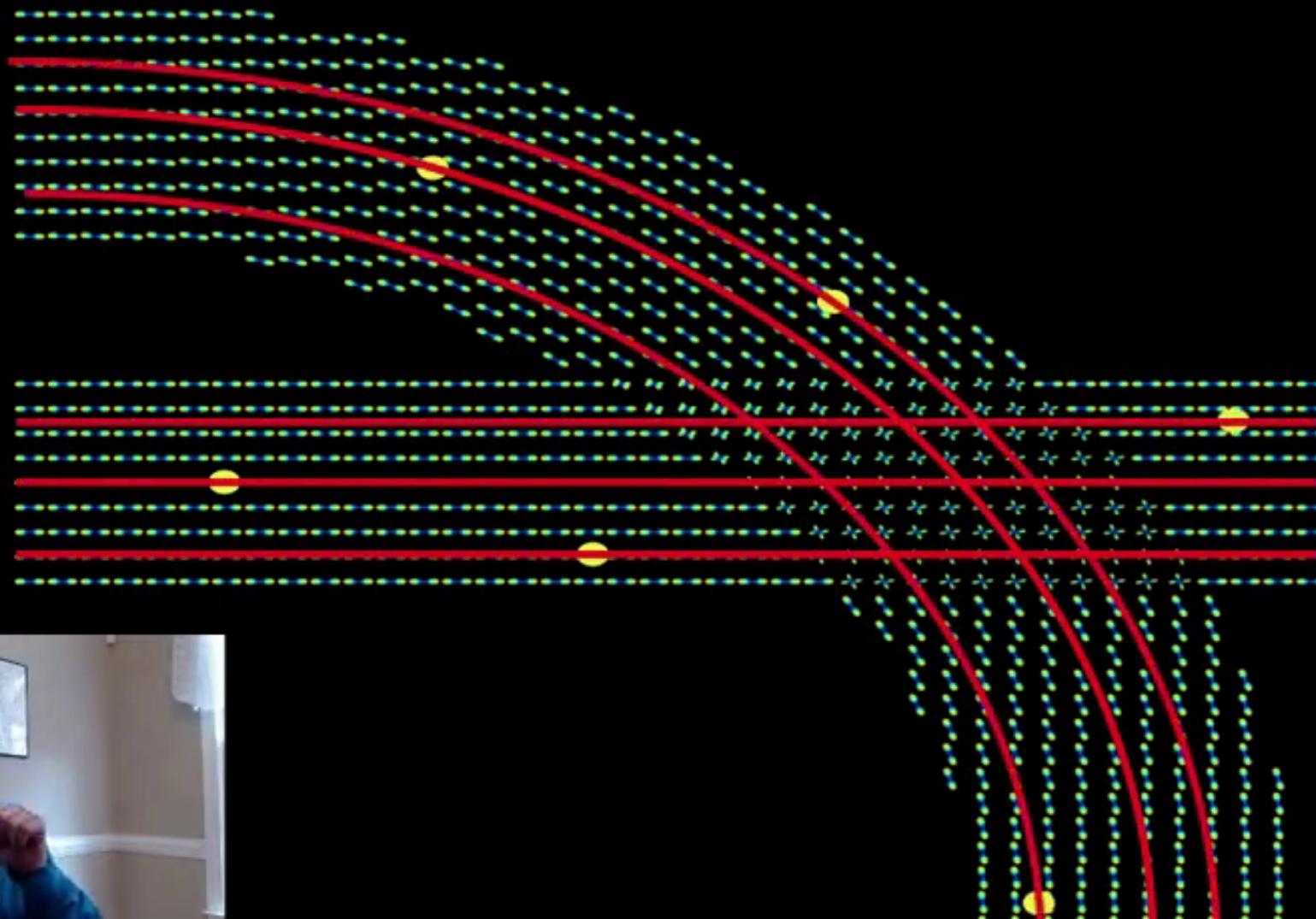
Hough Transform !!!

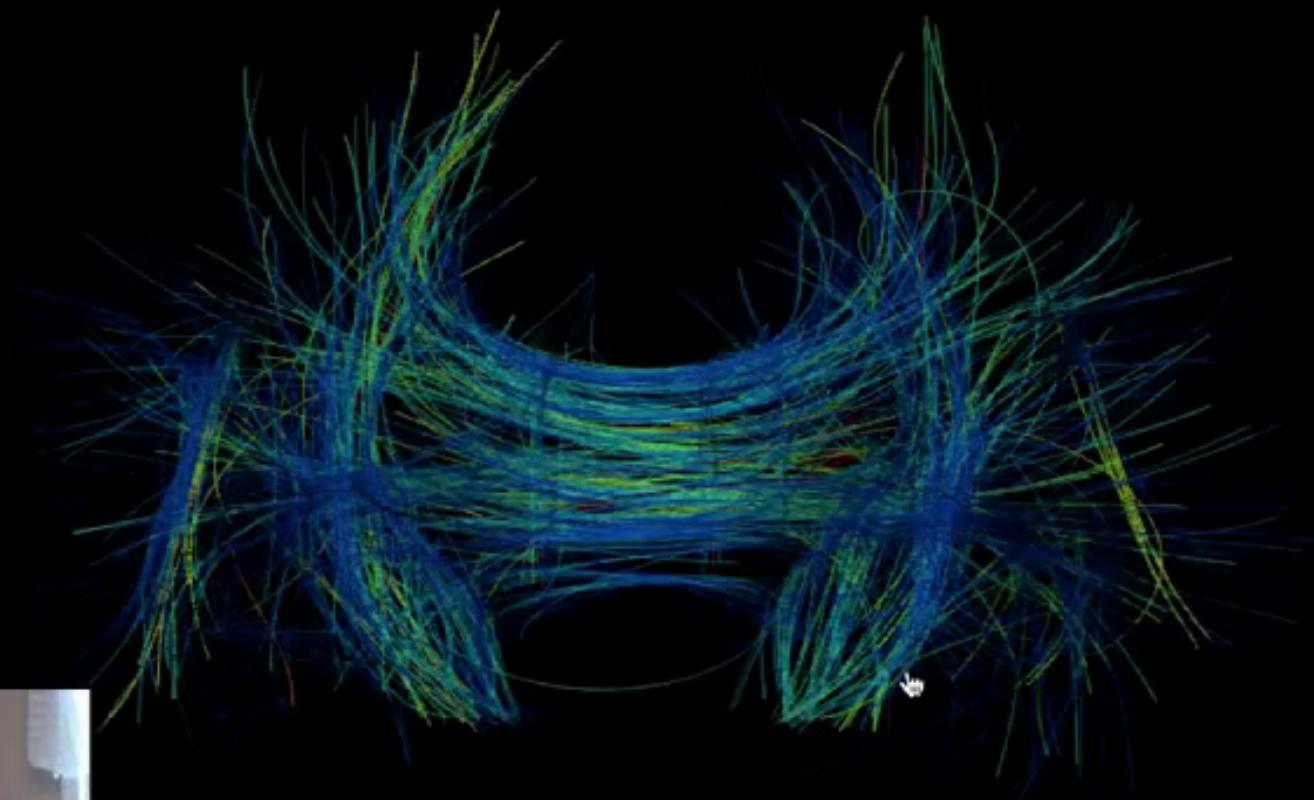


Hough Transform !!!



Hough Transform !!!





Brain Imaging: Deep Brain Stimulation

**Image and Video Processing: From Mars to
Hollywood with a Stop at the Hospital**

Guillermo Sapiro

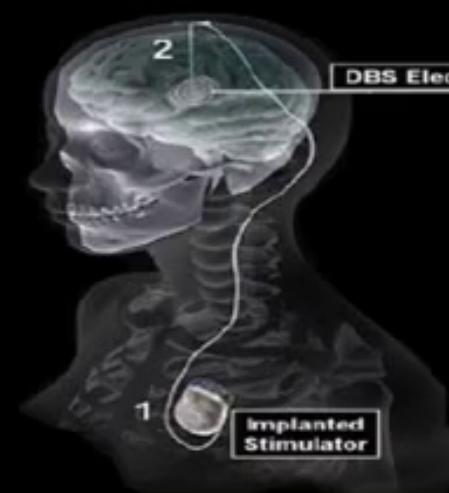


Thanks to Lenglet, Aganj, Harel, Duchin, SIS

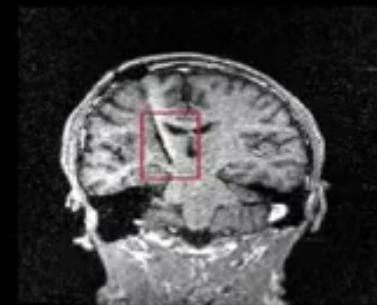




Deep Brain Stimulation (DBS)



Cameron McIntyre, Cleveland Clinic Foundation



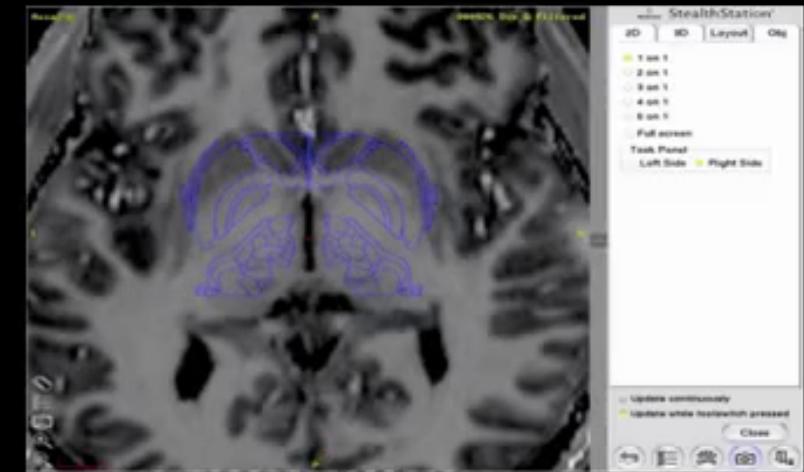
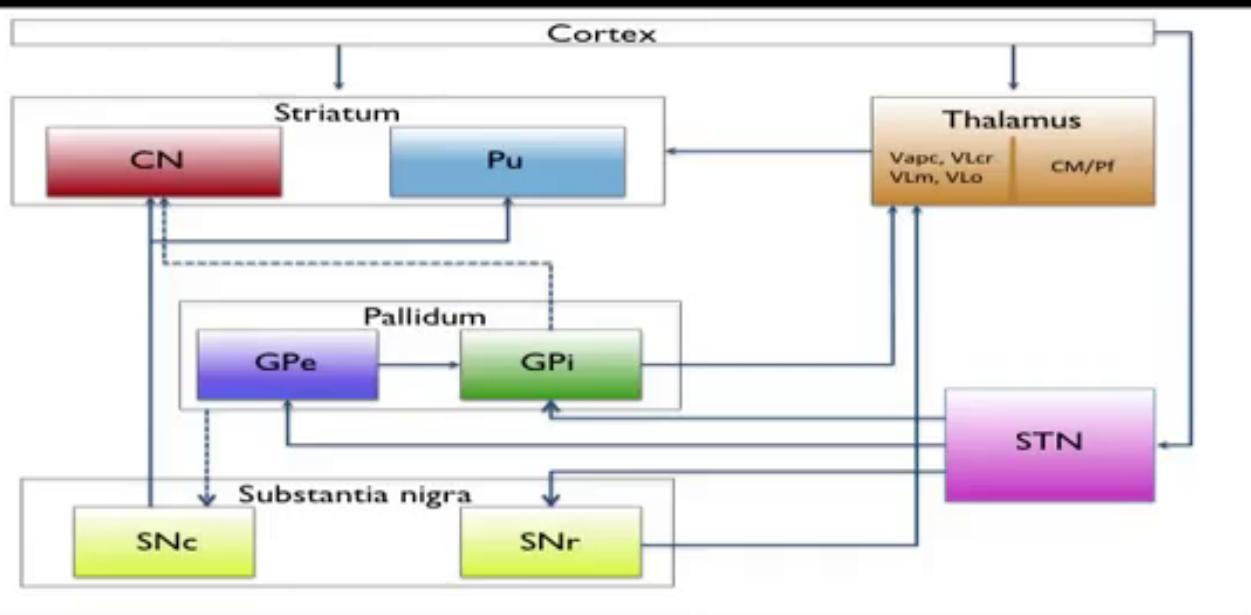
Successful DBS surgery is critically dependent on precise placement of DBS electrodes into target structures

Students: A good place to take a break if needed.

Duke
UNIVERSITY

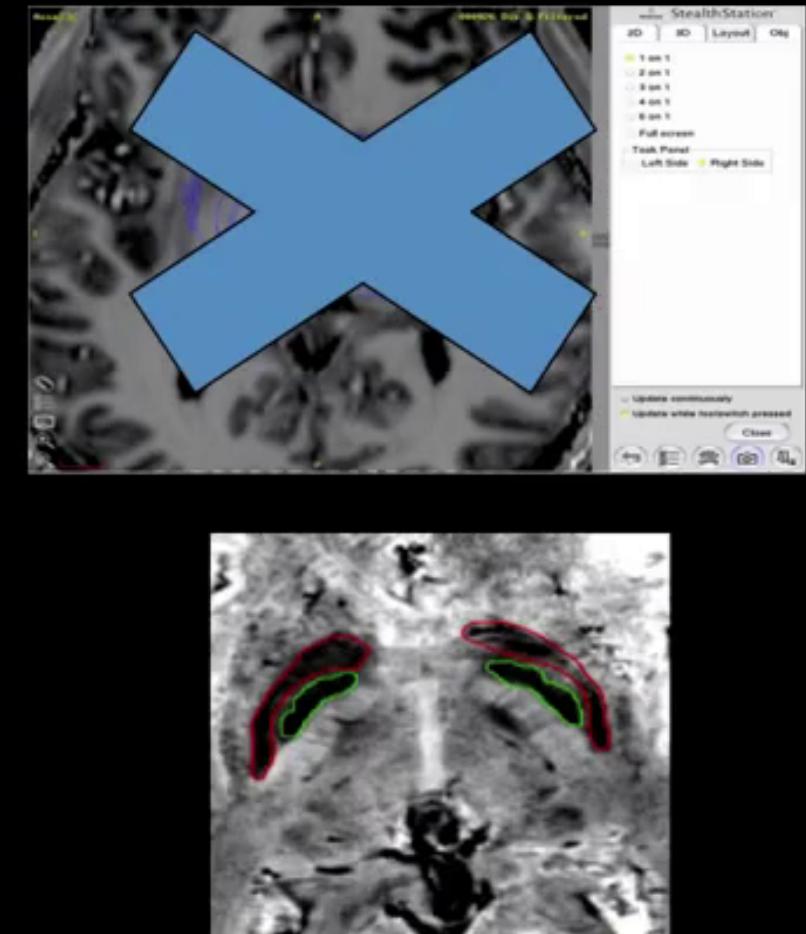
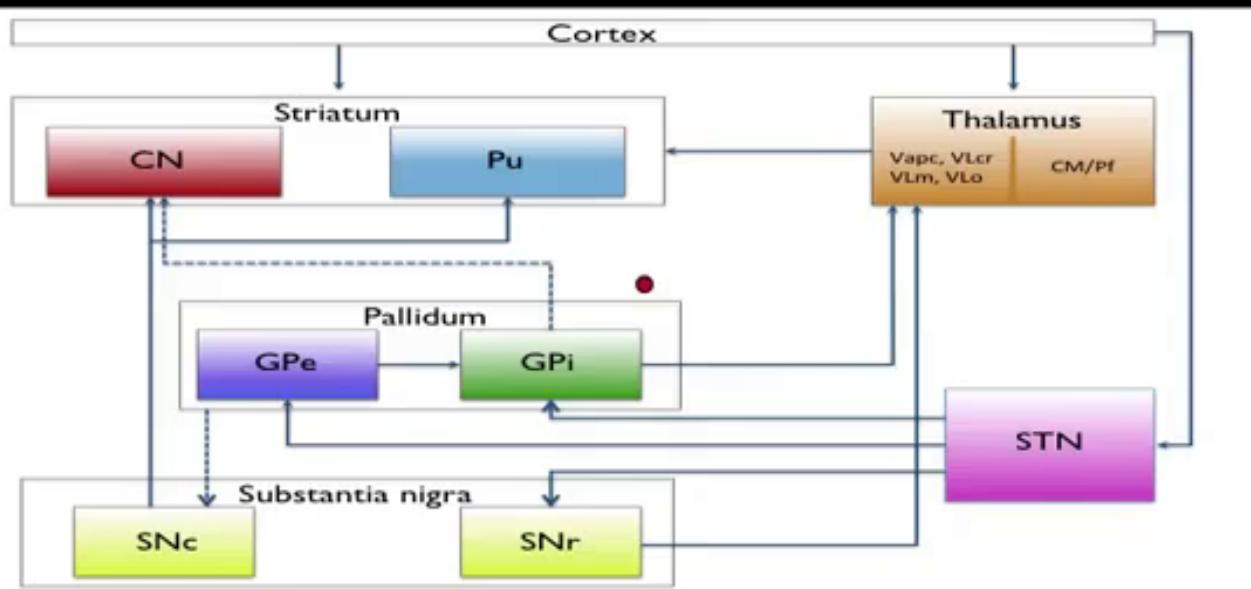


Brain Imaging and DBS



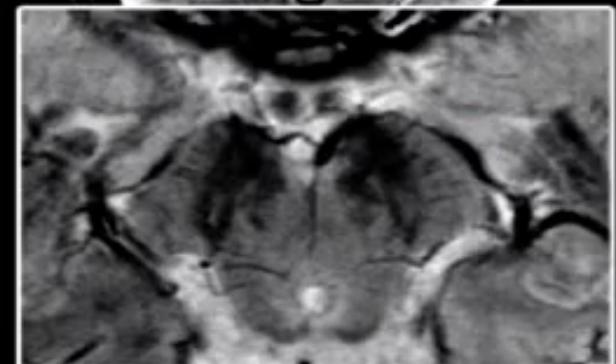
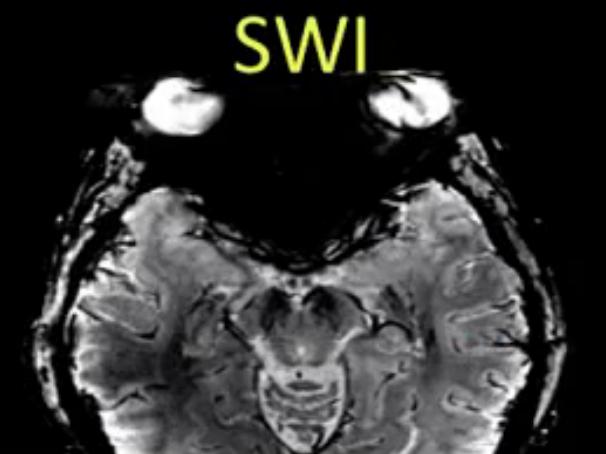
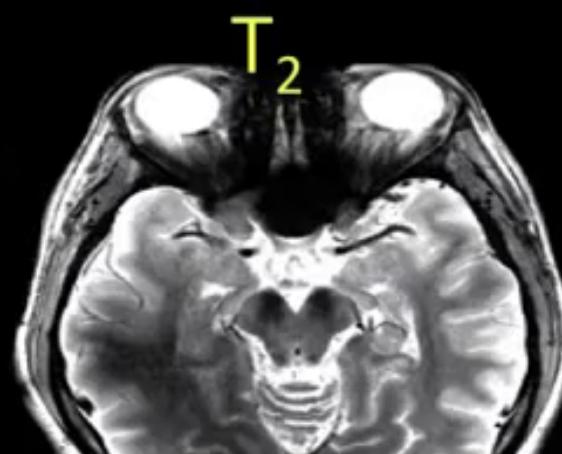
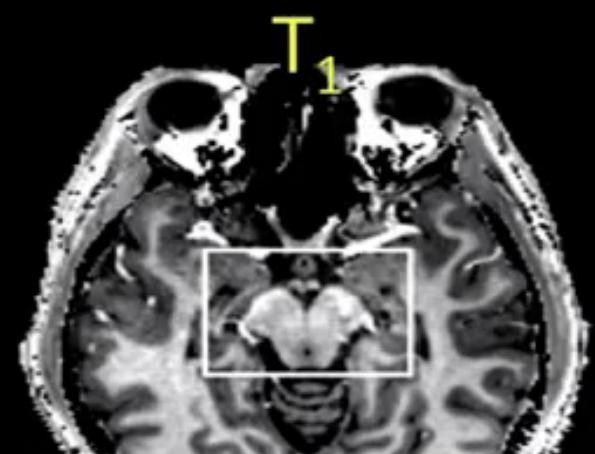


Brain Imaging and DBS



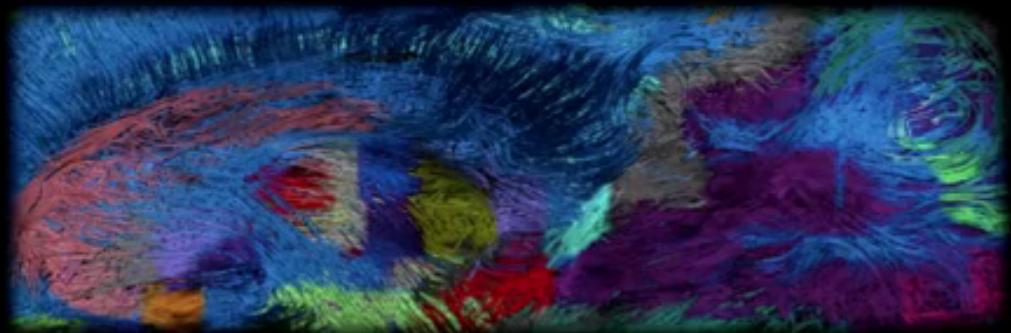
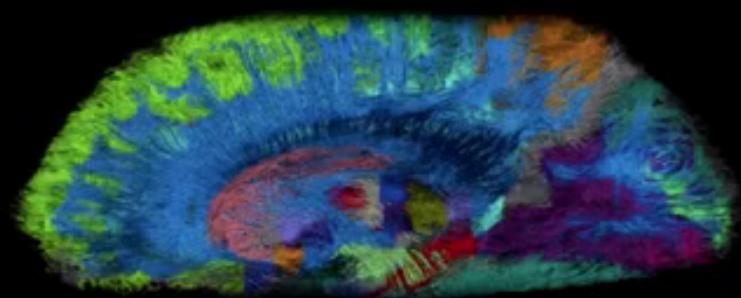
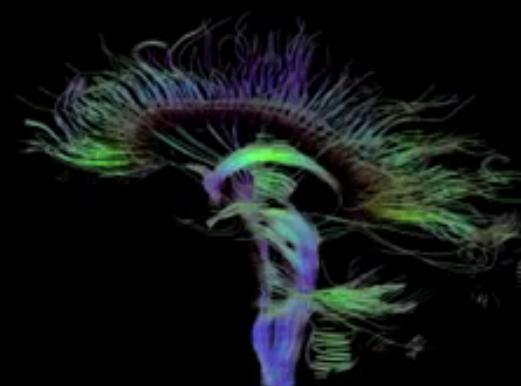
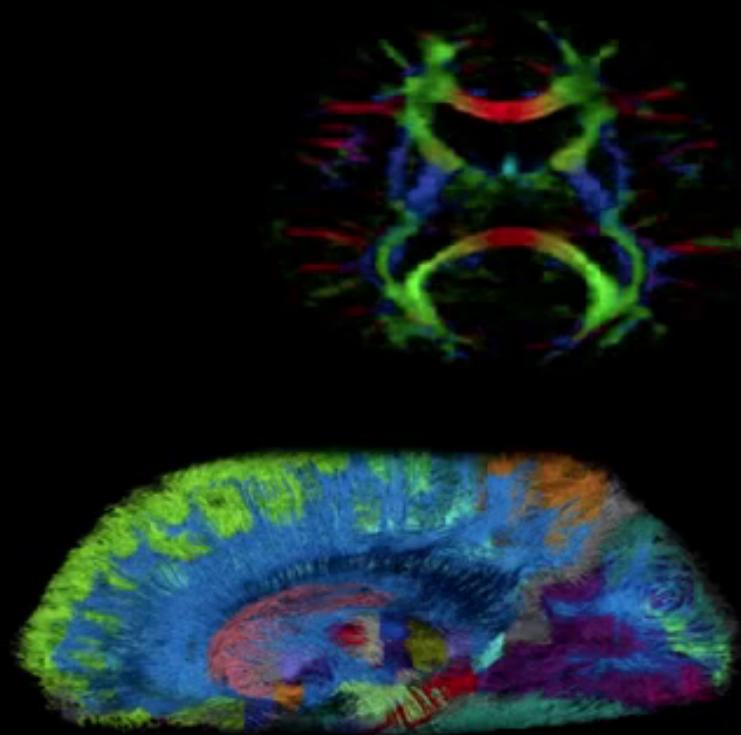


Brain Imaging and DBS





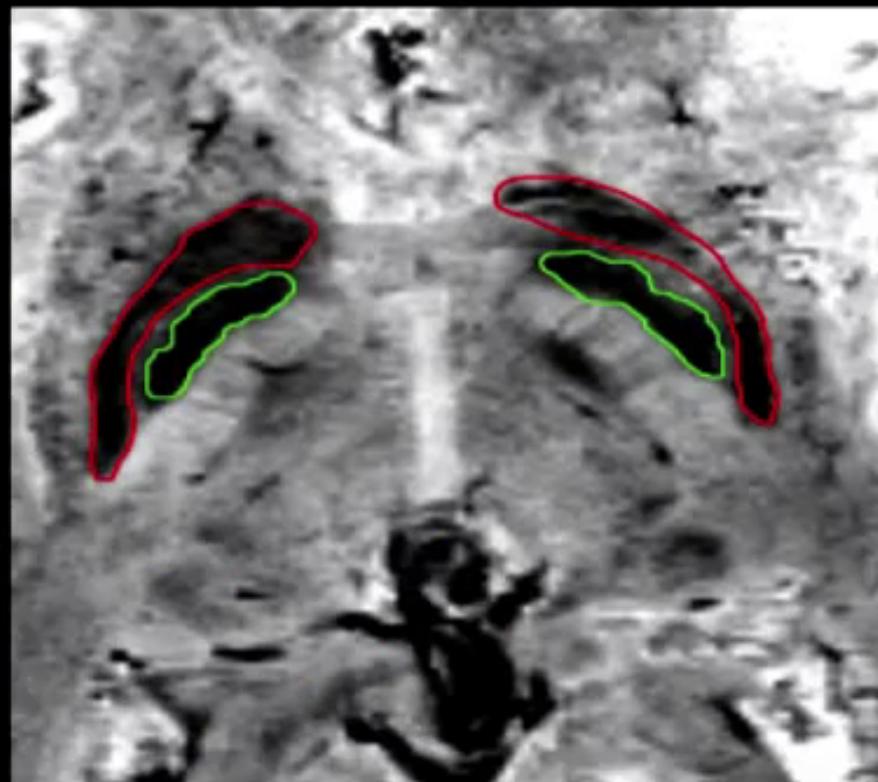
Brain Imaging and DBS



Human, 7T, $1.5 \times 1.5 \times 1.5 \text{ mm}^3$
Tractography – Paul Thompson, UCLA



Brain Imaging and DBS

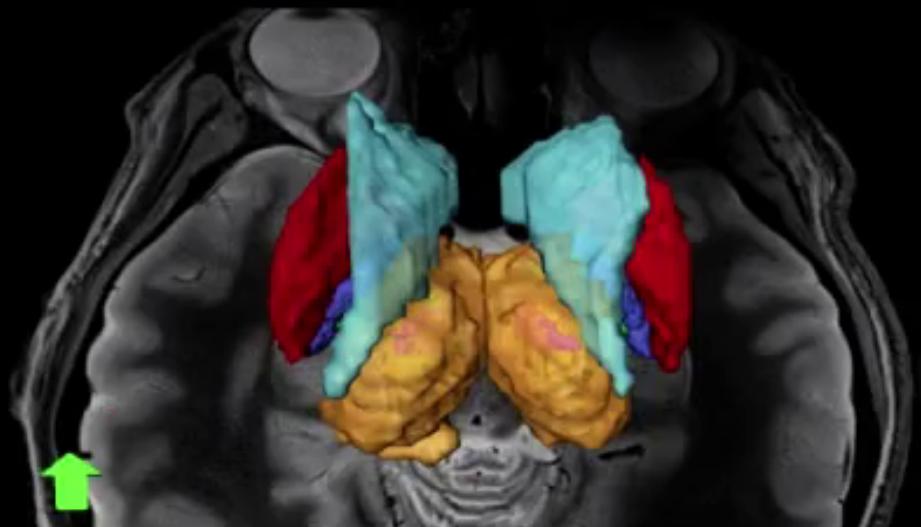


GP = Globus pallidus

GPi = DBS Target for Dystonia



Brain Imaging and DBS: ROI





Brain Imaging and DBS: DWI

