

Image Analysis (02502)

Advanced Topics

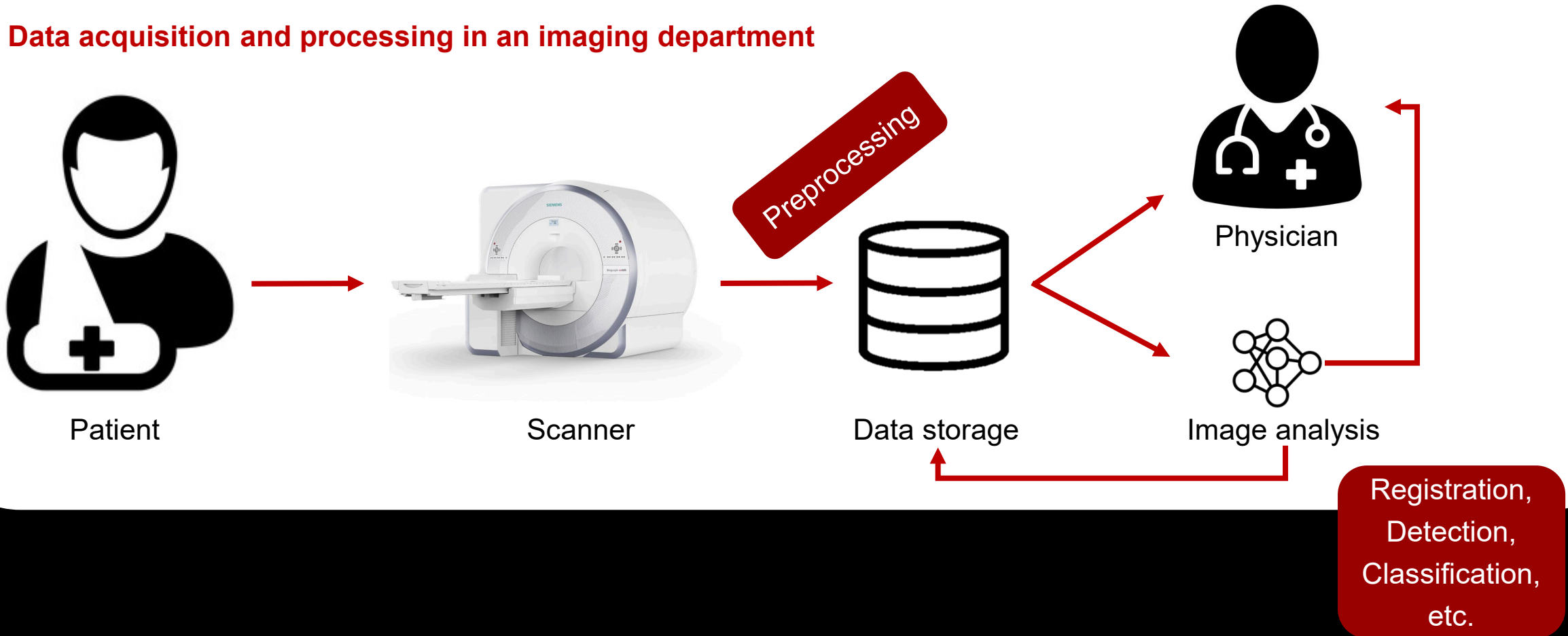
Claes Ladefoged, PhD

Claes N. Ladefoged

- MSc from Computer Science KU
- PhD in Medicine from SUND
- Head of AI Research at Rigshospitalet
- Associate Professor, DTU Compute

Overview

Data acquisition and processing in an imaging department

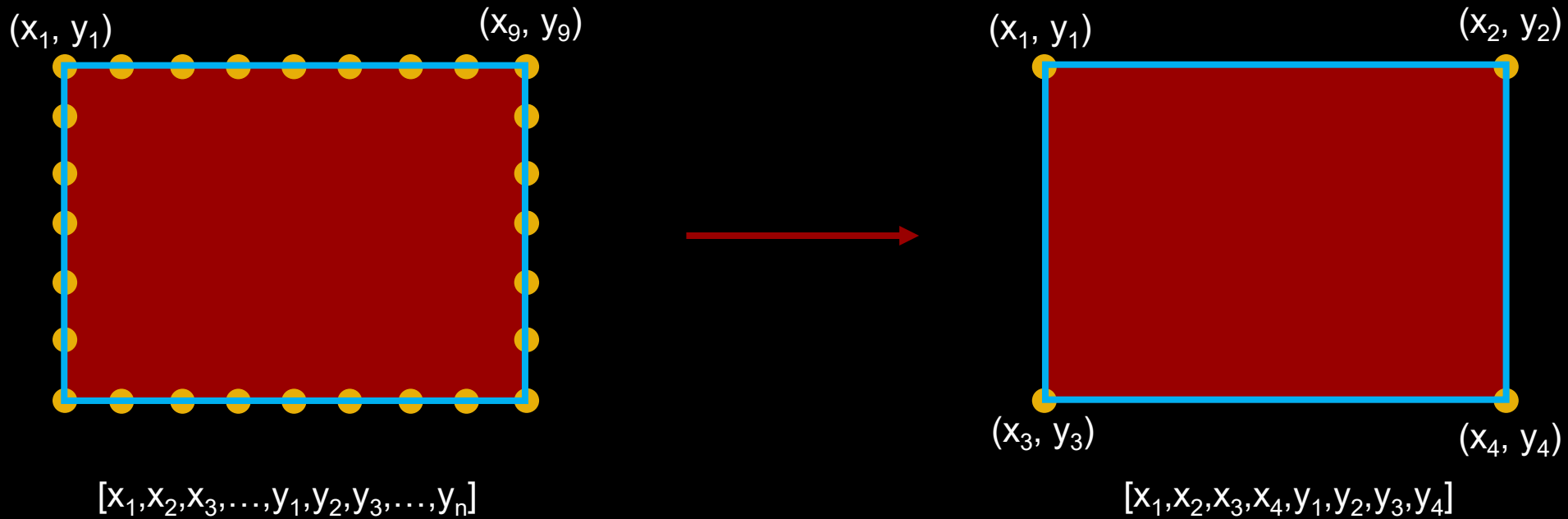


Preprocessing

- Data compression
- Intensity normalization
- Intensity Augmentation
- Intensity mapping
- Filtering

Data compression

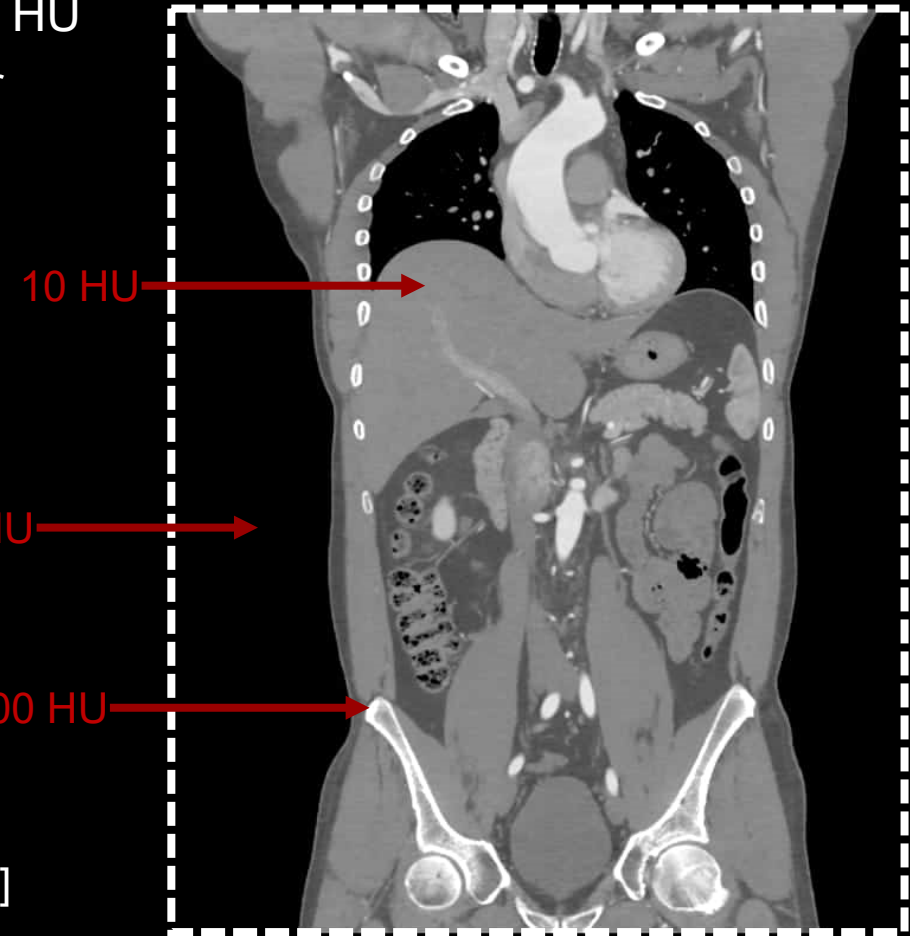
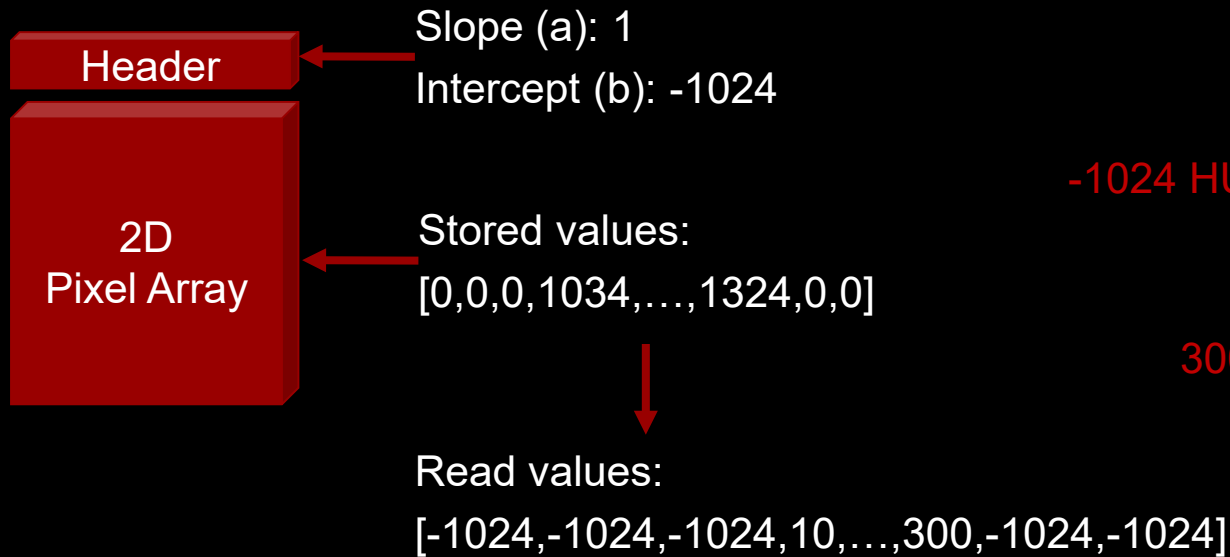
- Representation of outlines



Data compression

- CT values are usually defined in [-1024;3071] HU
- Values are usually stored as unsigned integer
- Large part of the volume is air (-1024 HU)

$$F(x) = ax + b$$



Quiz 1

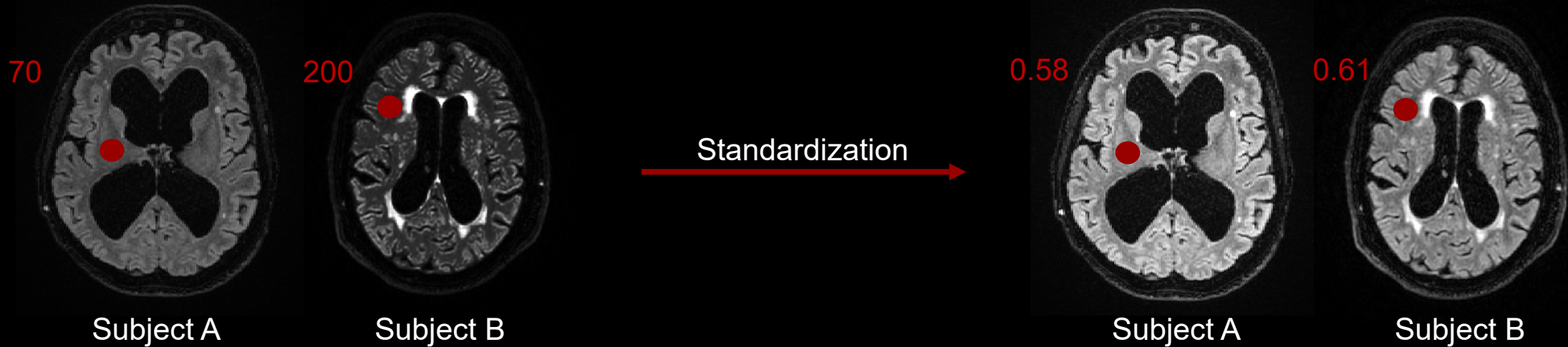
- An image containing values ranging from 0 to 52,427 needs to be stored in DICOM format
- The DICOM file has to be in the type SHORT (max value = 32,767)
- What can the slope and intercept be?
 - Slope 1.4 and intercept 1
 - Slope 1.6 and intercept 0
 - Slope 1 and intercept -19,660

menti.com

Code: **7419 1151**

Intensity normalization

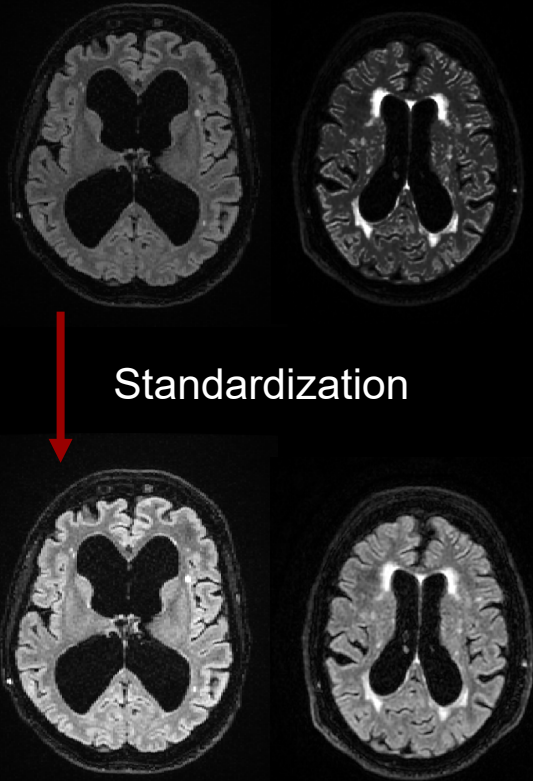
- Conventional MRI intensities (T1-w, T2-w, PD, FLAIR) are acquired in arbitrary units



Quantification analysis:
Incomparable results

Quantification analysis:
Comparable results

Intensity normalization



Some available mapping functions:

- Min-max scaling

$$g(x, y) = \frac{f(x, y) - v_{min}}{v_{max} - v_{min}}$$

- Histogram stretching

$$g(x, y) = \frac{v_{max,d} - v_{min,d}}{v_{max} - v_{min}} (f(x, y) - v_{min}) + v_{min,d}$$

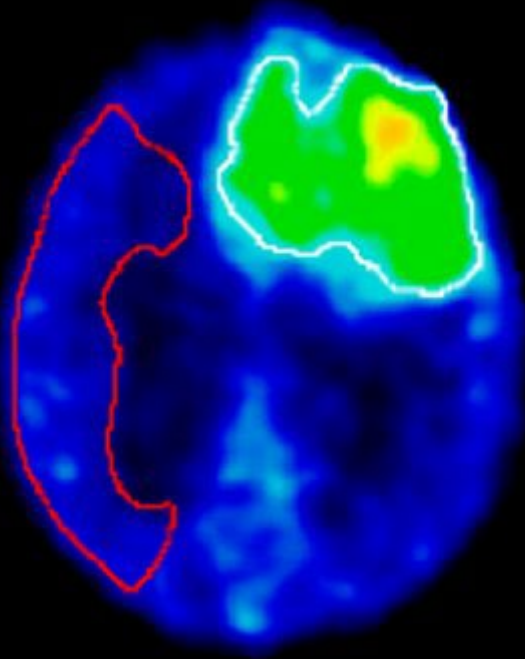
- Z-normalization

$$g(x, y) = \frac{f(x, y) - \mu}{\sigma}$$

Be aware when high intensity areas are present!

Z-normalization is the de-facto standard for most MRI-based preprocessing
 What about images with non-arbitrary units (CT, PET)?

Intensity normalization



Normalize relative to a reference region before scaling

Examples:

- Background region in brain
- Liver region in whole-body imaging

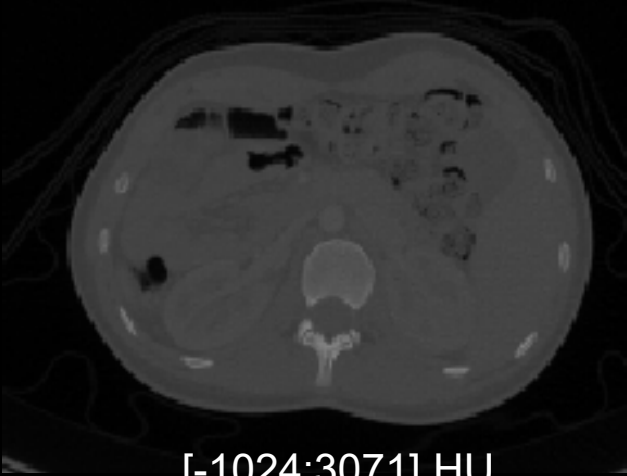
Intensity mapping

E.g. by histogram stretching or intensity rescaling:

Each image is mapped from v_{min} and v_{max} to $v_{min,d}$ and $v_{max,d}$ (often 0-255) using:

$$g(x, y) = \frac{f(x, y) - v_{min}}{v_{max} - v_{min}} * (v_{max,d} - v_{min,d}) + v_{min,d}$$

followed by clamping values outside the range



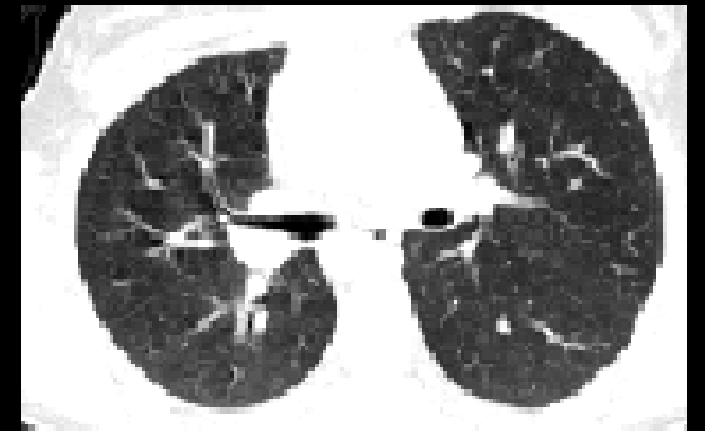
[-1024;3071] HU



[-150;250] HU

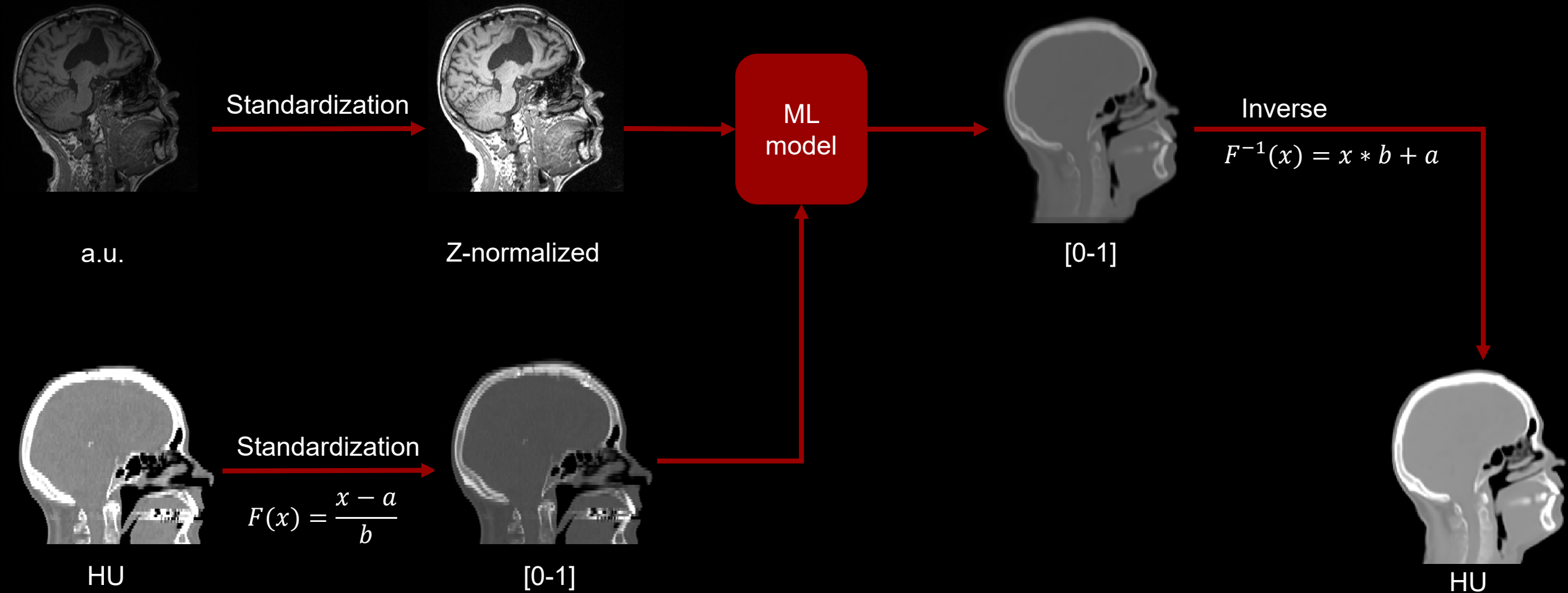


[-150;250] HU

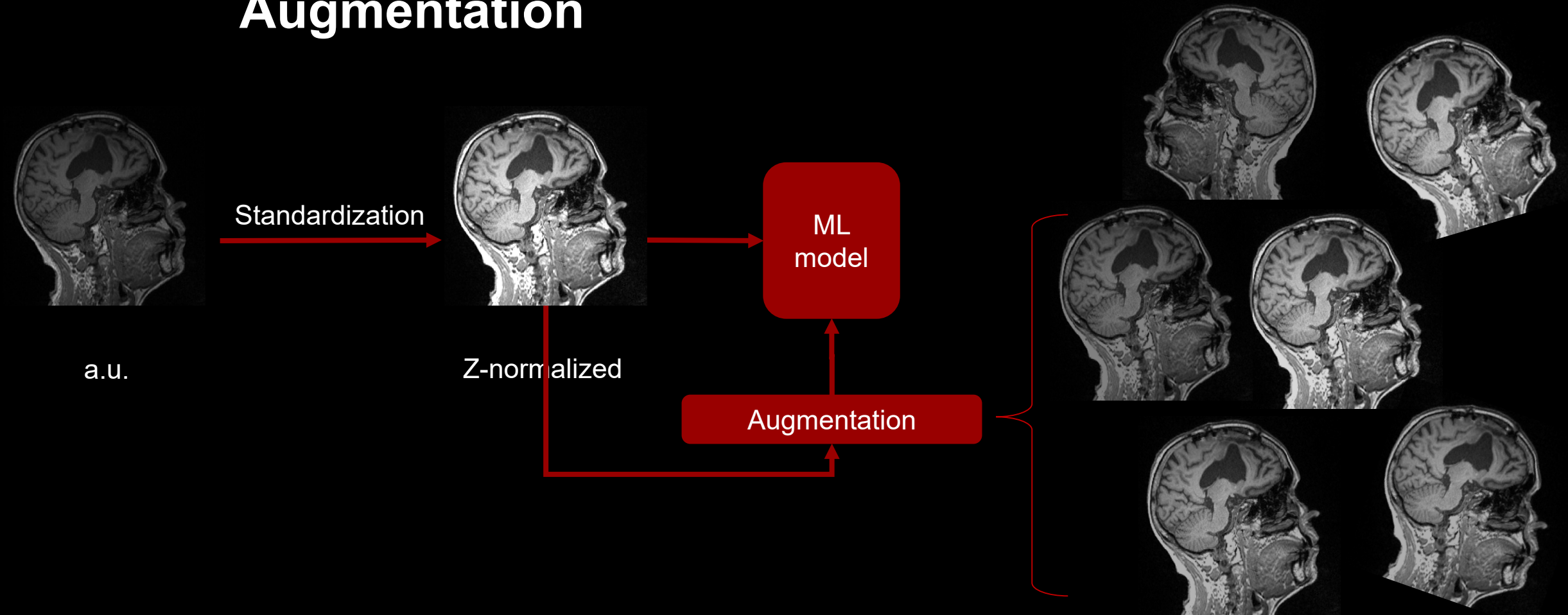


[-1000;0] HU

Intensity normalization



Augmentation



Quiz 2

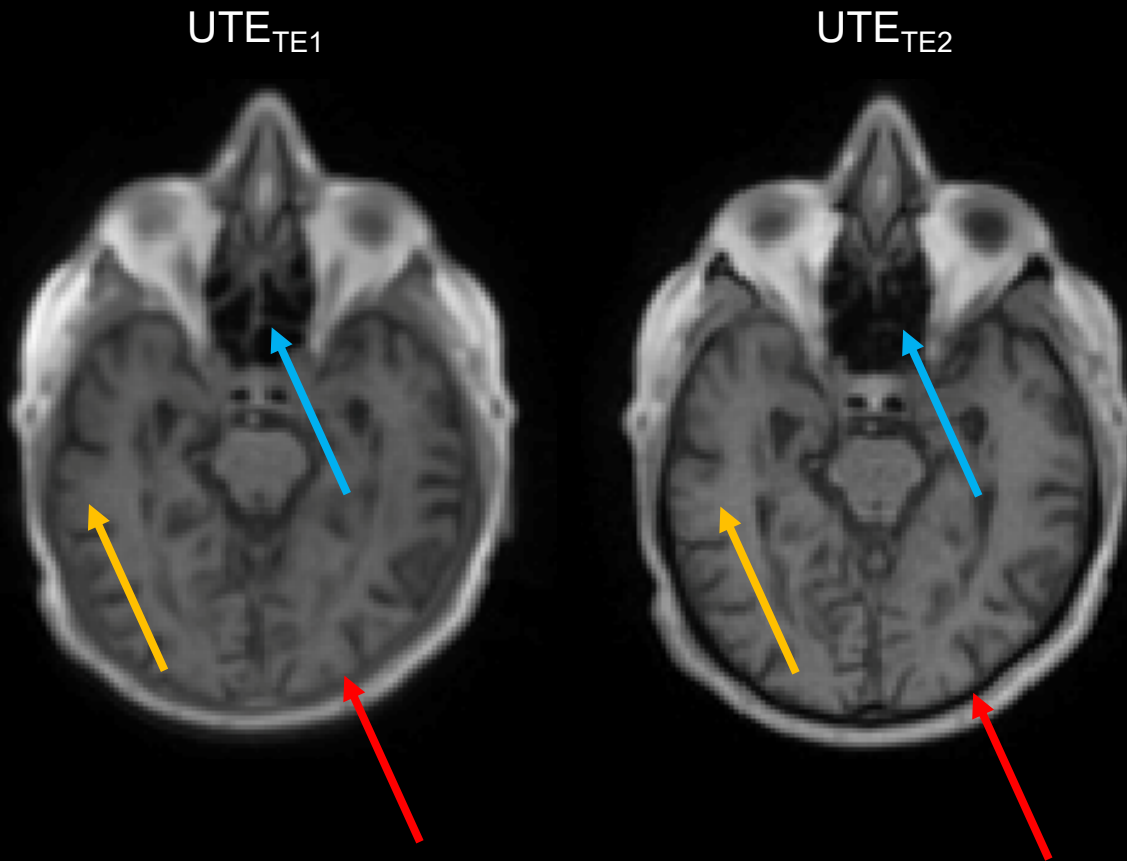
- A model is trained to predict the perceived age of a patients' brain given an MRI
- The model was trained with data containing ages of 18 to 99, so was scaled using:

$$g(x, y) = \frac{f(x, y) - v_{min}}{v_{max} - v_{min}} * (v_{max,d} - v_{min,d}) + v_{min,d}$$

where $(v_{min}, v_{max}) = (18, 99)$ and $(v_{min,d}, v_{max,d}) = (0, 1)$

- The model predict 0.78 for a given MRI. What is the predicted age (in years) of the patient?
 - 63
 - 70
 - 81
 - 95

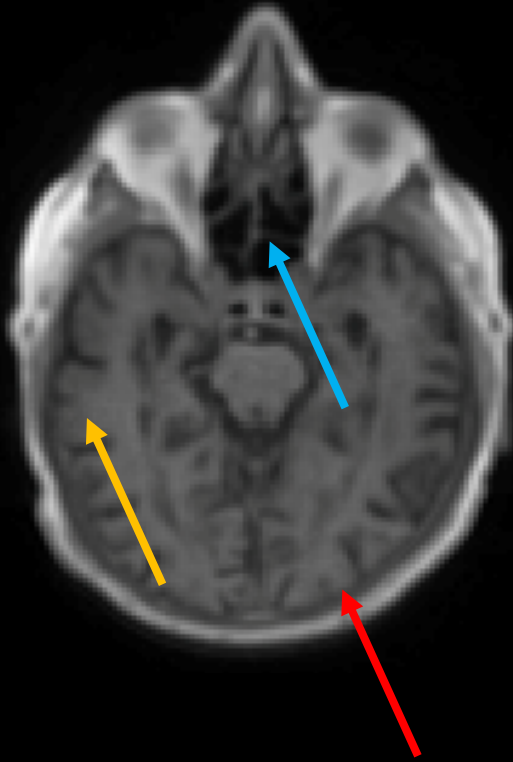
Intensity mapping



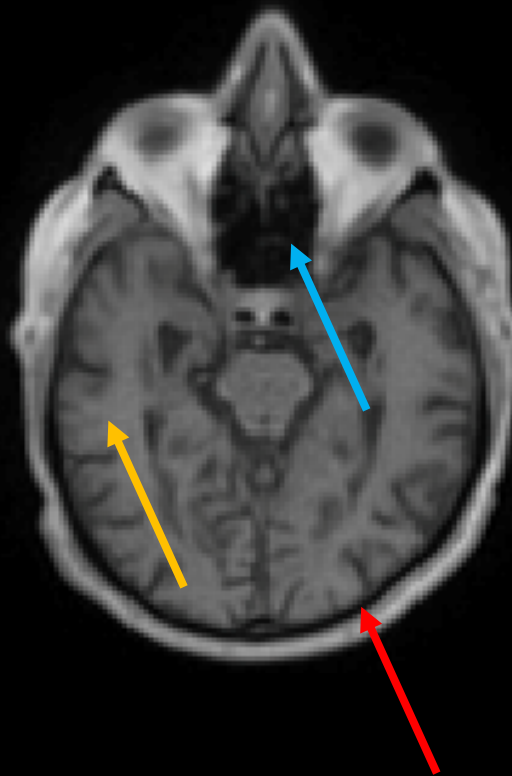
- Two MR images acquired with different echo times $TE1 \ll TE2$
- Different intensities are expected in **bone** but not in **air** and **tissue**

Intensity mapping

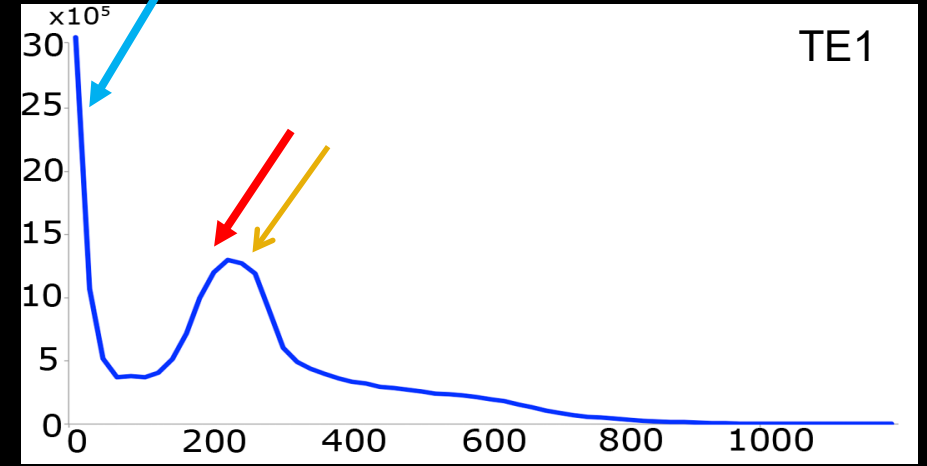
UTE_{TE1}



UTE_{TE2}

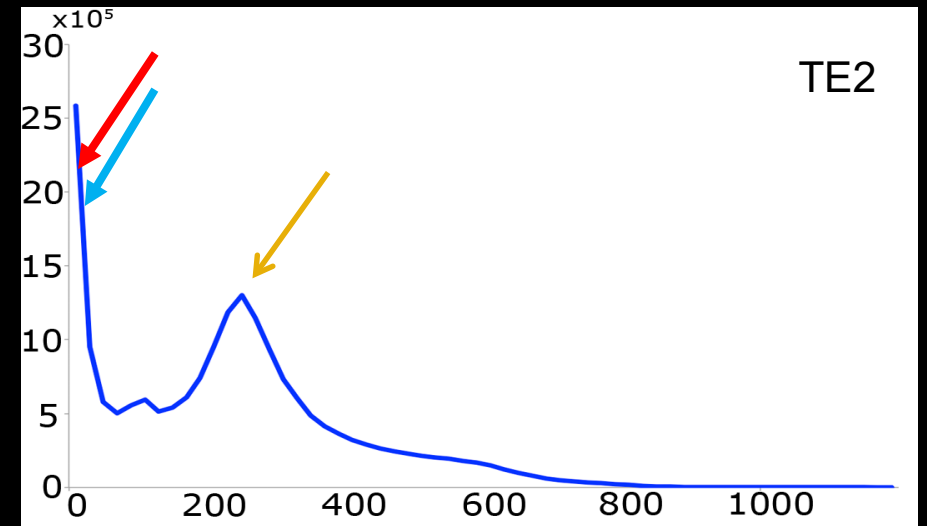


Frequency



MR intensity

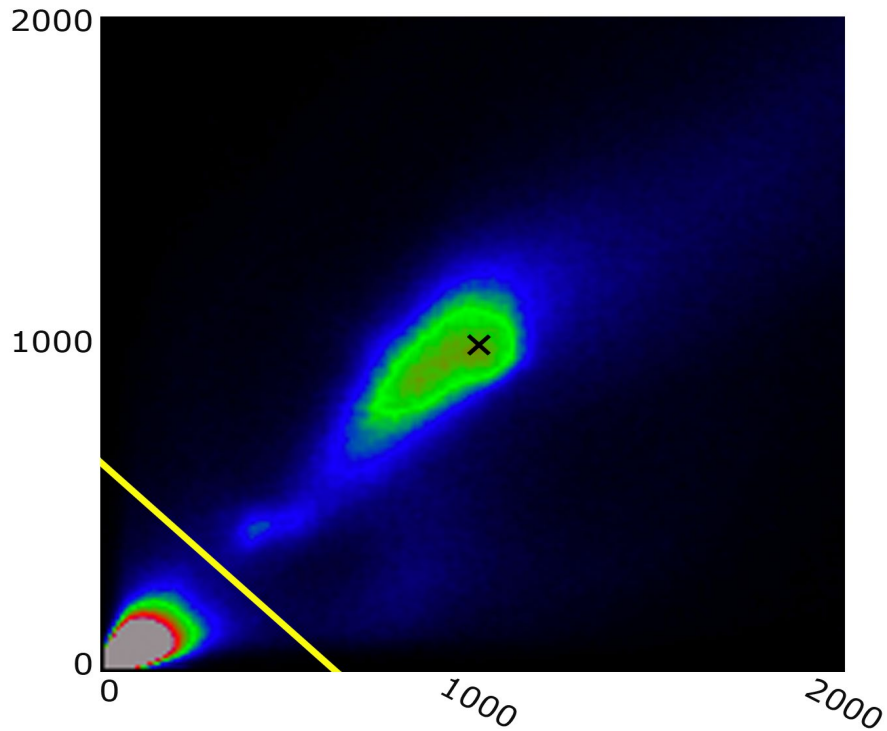
Frequency



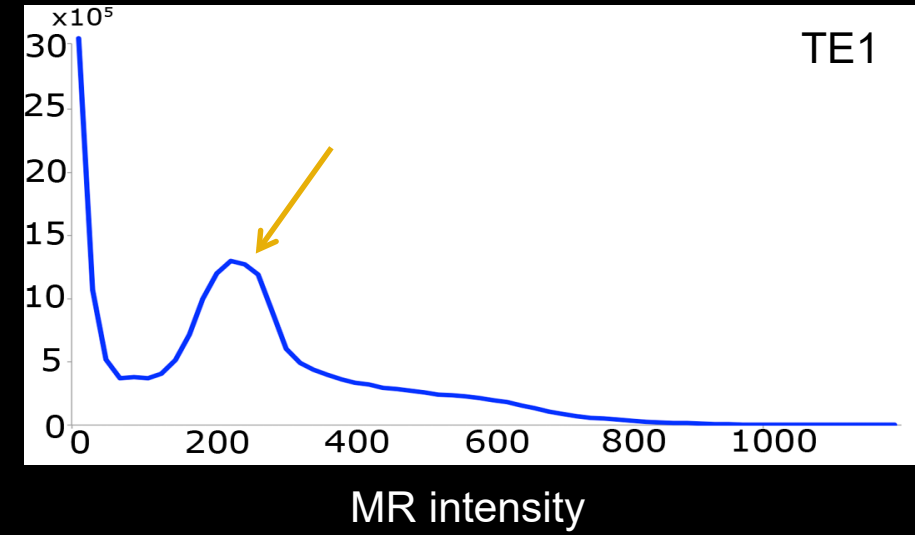
MR intensity

Intensity mapping

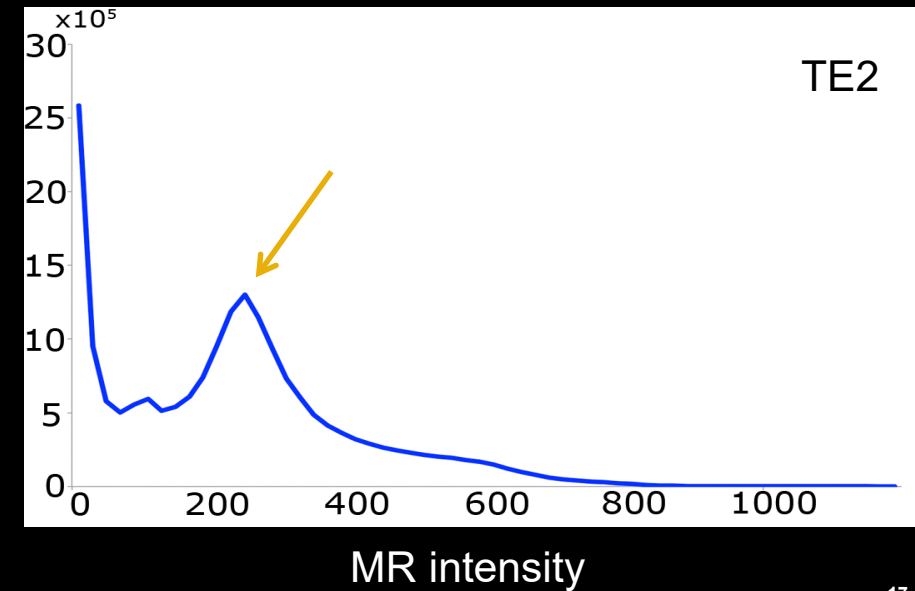
Normalized Joint histogram



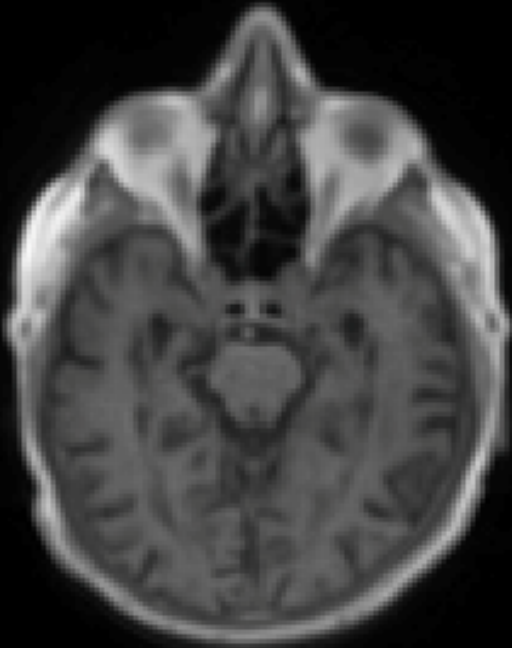
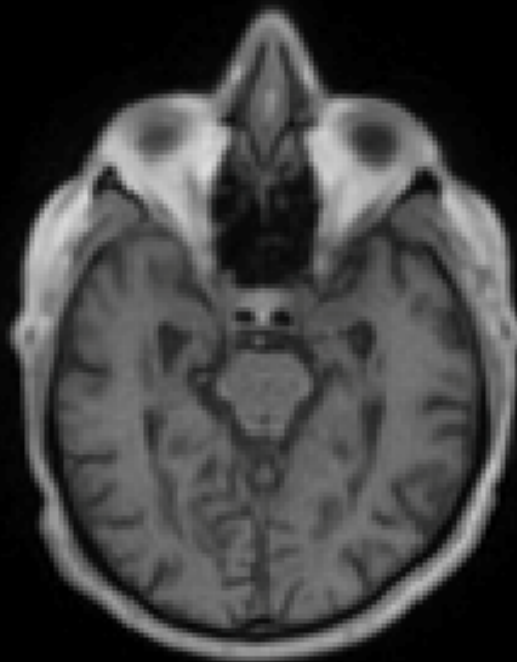
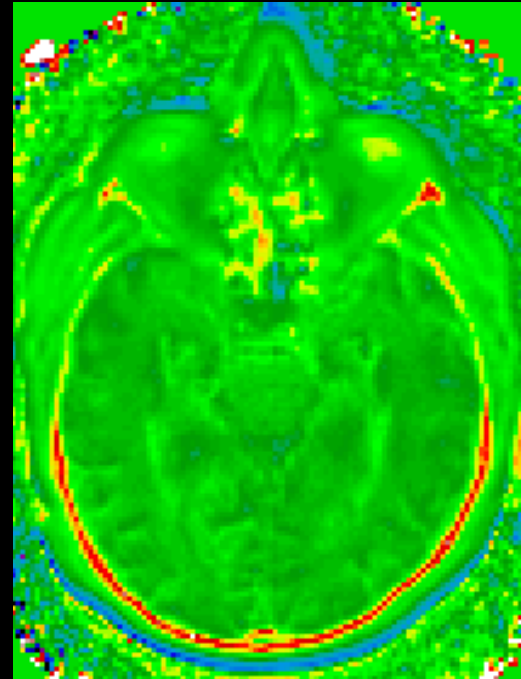
Frequency



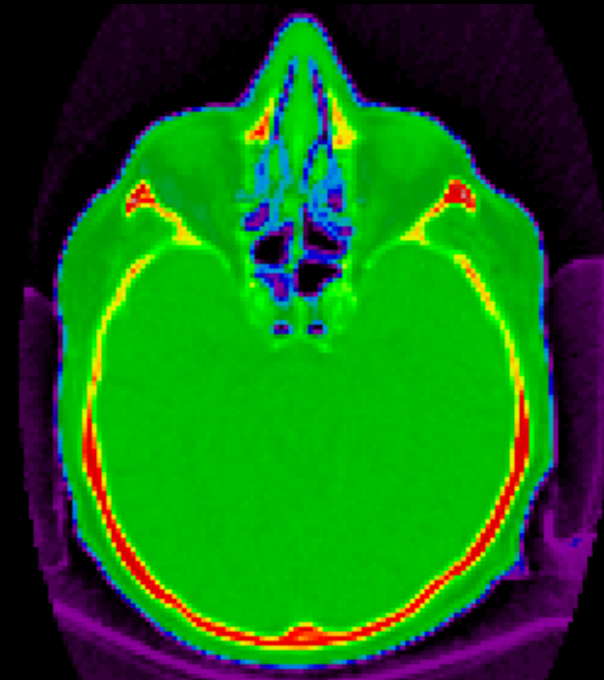
Frequency



Intensity mapping

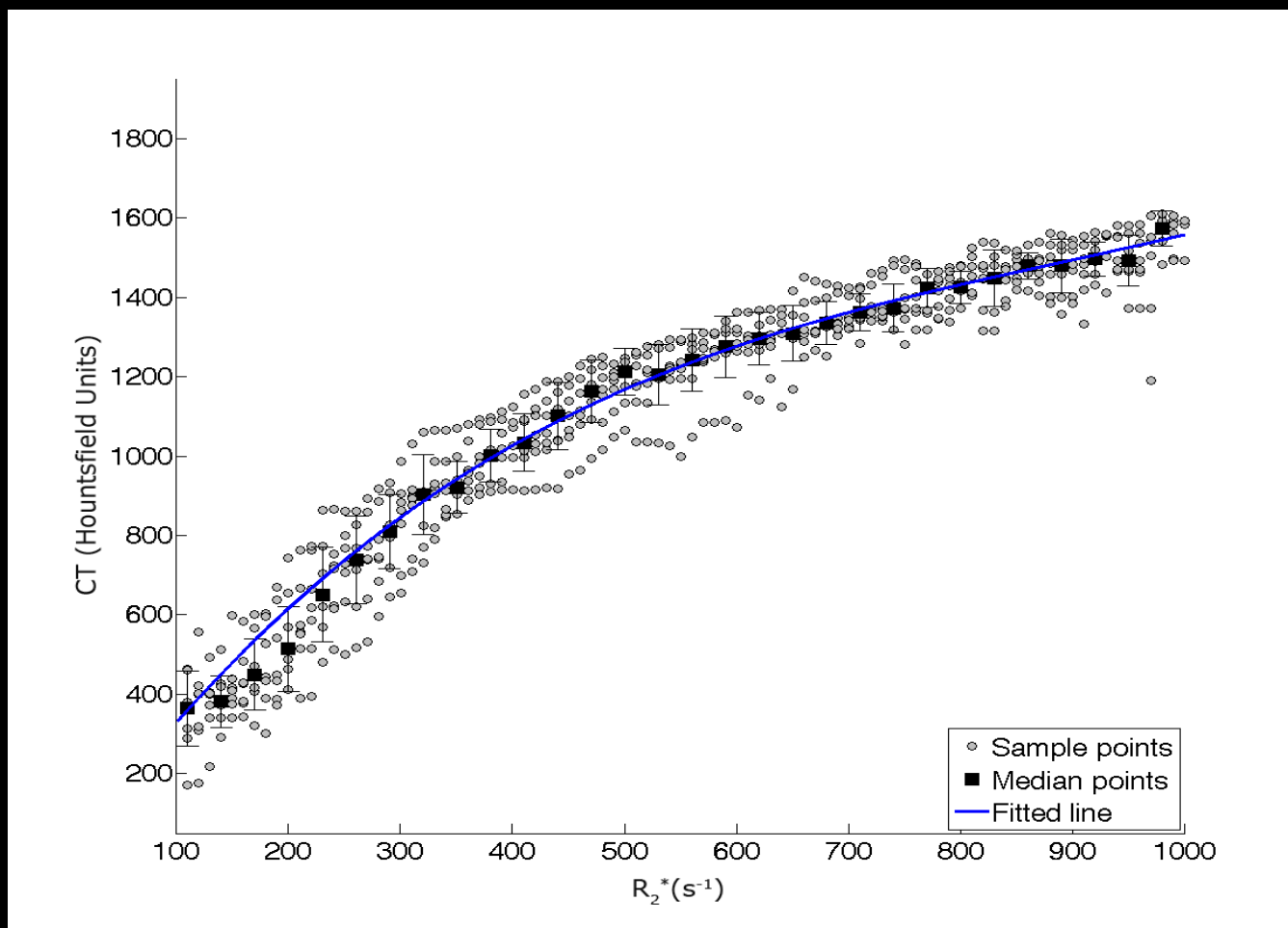
 UTE_{TE1}  UTE_{TE2}  R_2^* 

CT

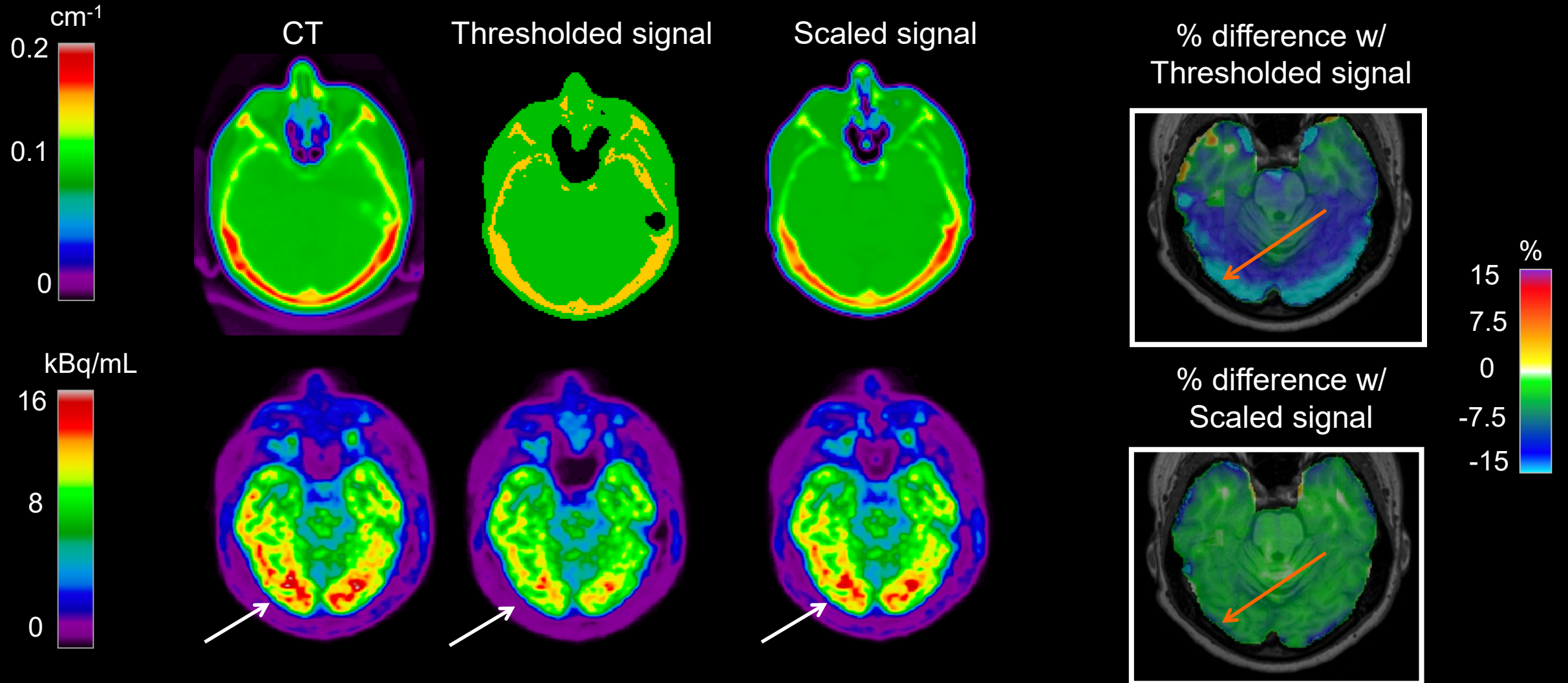


$$R_2^* = \frac{\ln(UTE_{TE1}) - \ln(UTE_{TE2})}{TE2 - TE1}$$

Intensity mapping



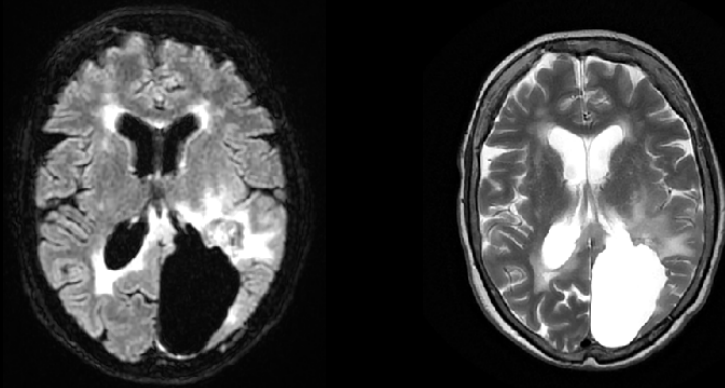
Intensity mapping



Registration

- Interpolation
- Intra subject registration
 - Same session
 - Between sessions
- Inter subject registration

Interpolation

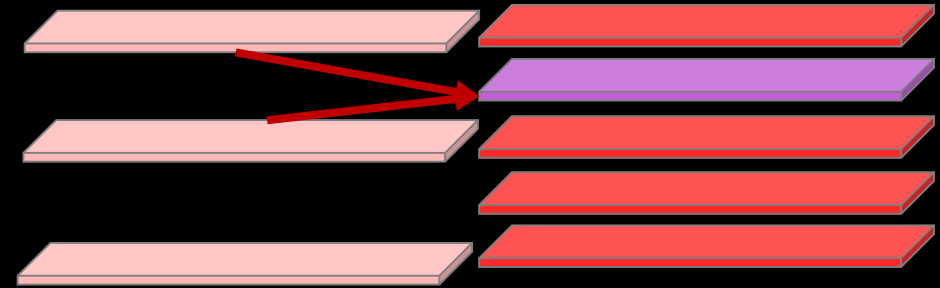
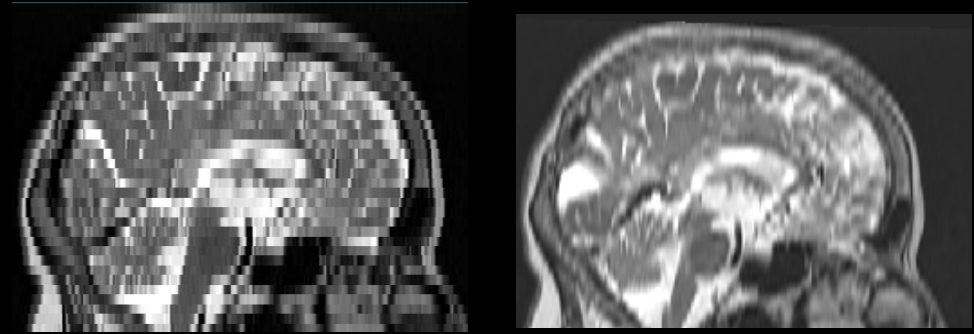


0.5x0.5x0.6 mm³

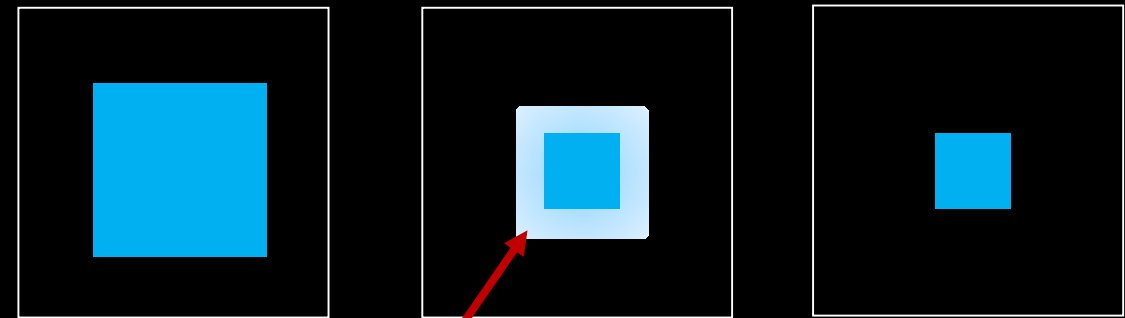
0.5x0.5x4.4 mm³



Image interpolation → Trilinear (or similar)



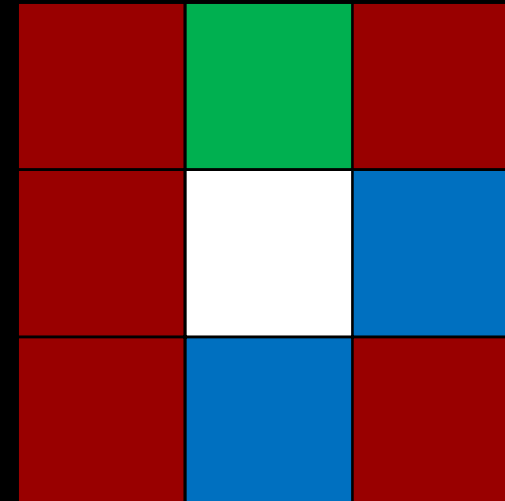
Label interpolation → Nearest Neighbour



Nearest neighbour ensures integer (e.g. 0 and 1) values

Quiz 3

- In a 4-connectivity setting, what would the color of the white center pixel be assigned when using nearest neighbour interpolation?
 - Green
 - Blue
 - Red



Registration

- Intra subject

Different transformations:

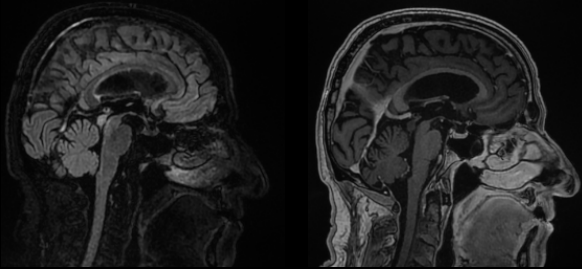
Translation

Rotation

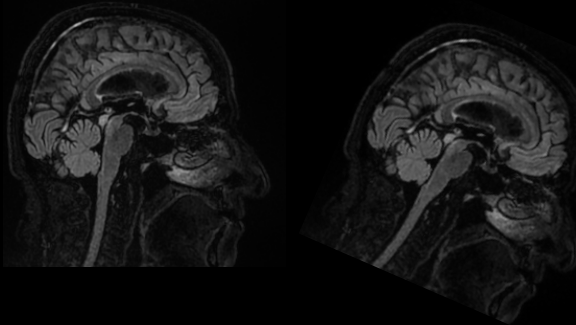
Scaling

Sheering

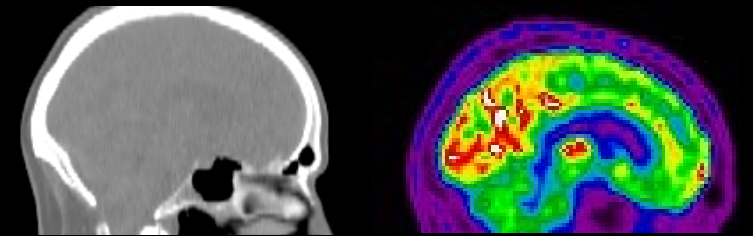
Between two similar modalities



Between two timepoints



Between two different modalities

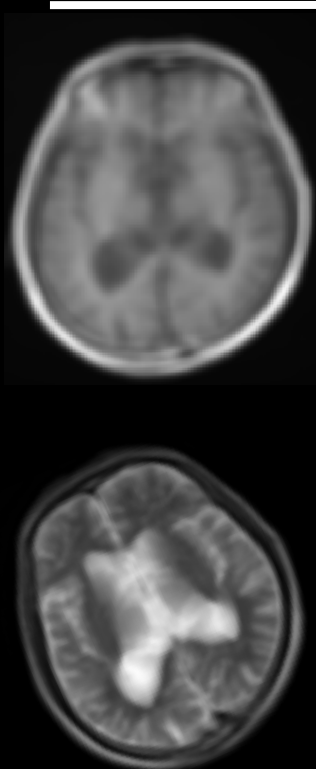
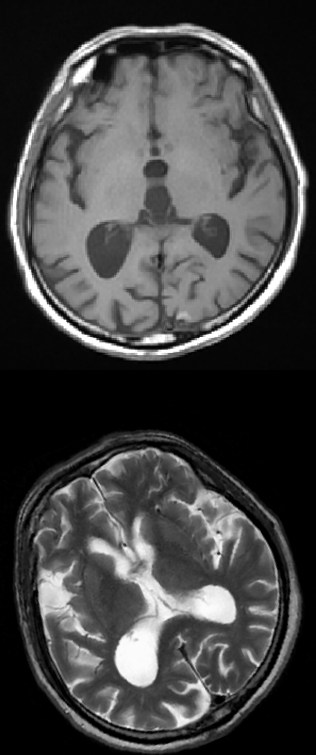


Translation and rotation are used for intra subject registration
Scaling mainly used for inter subject registration

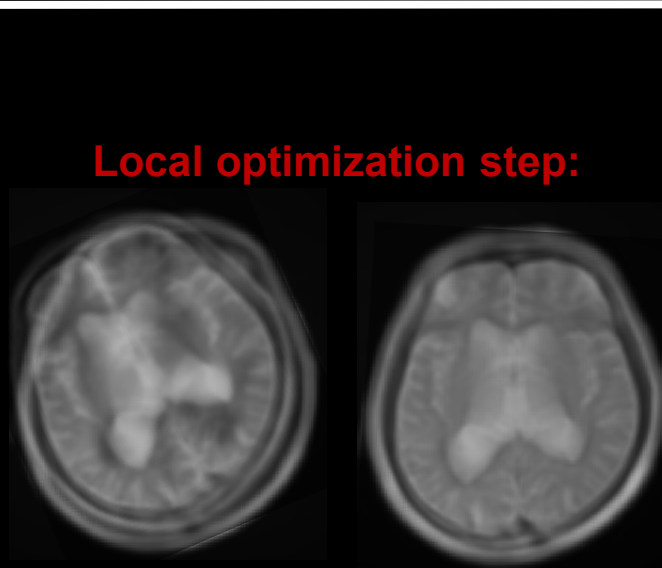
Registration

Global step:

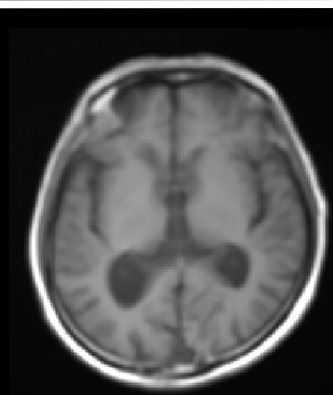
Search for overlap at low-to-high resolution



8mm



Course search grid to find optimal translation and rotation



4mm

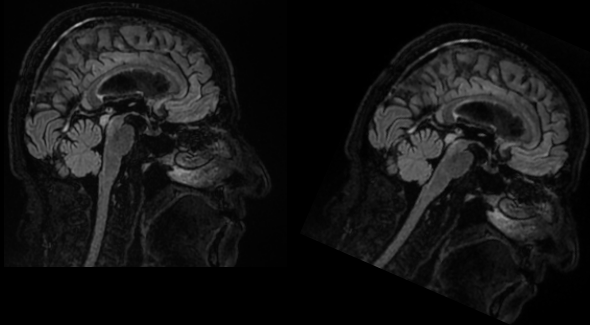


2mm

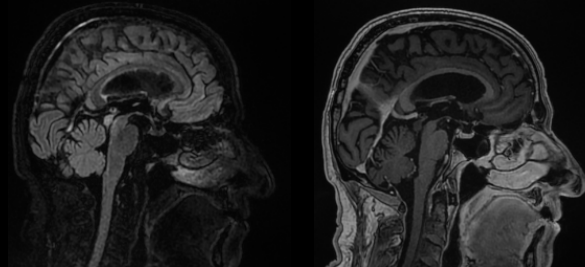
Registration

Similar modality cost function:
 Least squares
 Normalized correlation

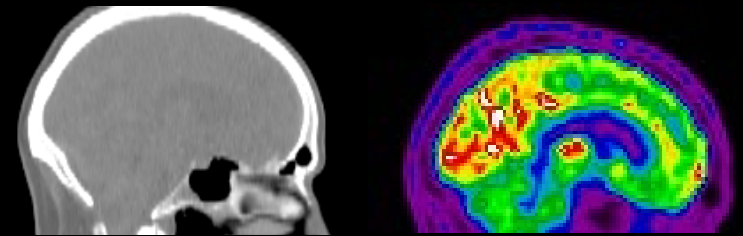
Between two timepoints



Between two similar modalities



Between two different modalities



Before registration

Target Moving

Brain extraction

Target Moving

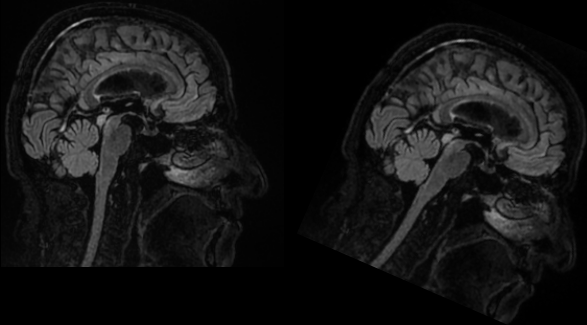
After registration

Target Moving

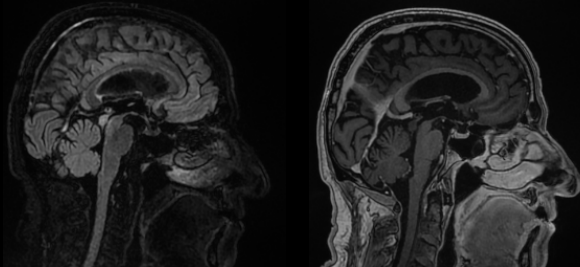
Registration

Different modality
cost-function:
Mutual information

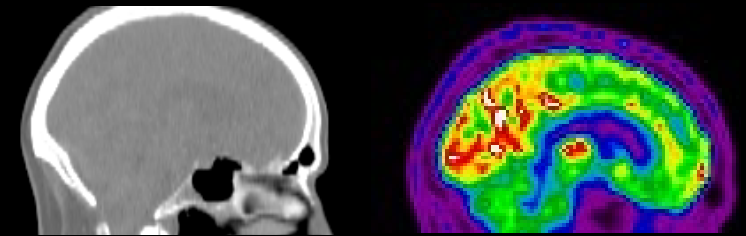
Between two timepoints



Between two similar modalities

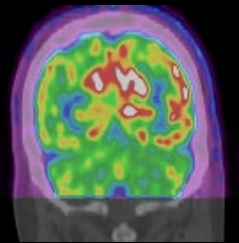
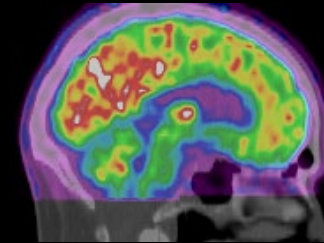
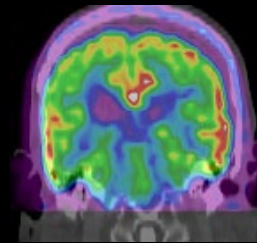
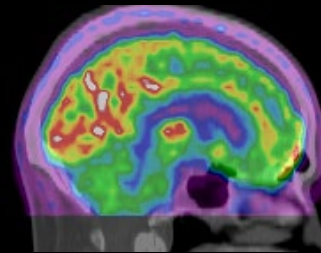


Between two different modalities



Before registration

After registration



Sagittal

Coronal

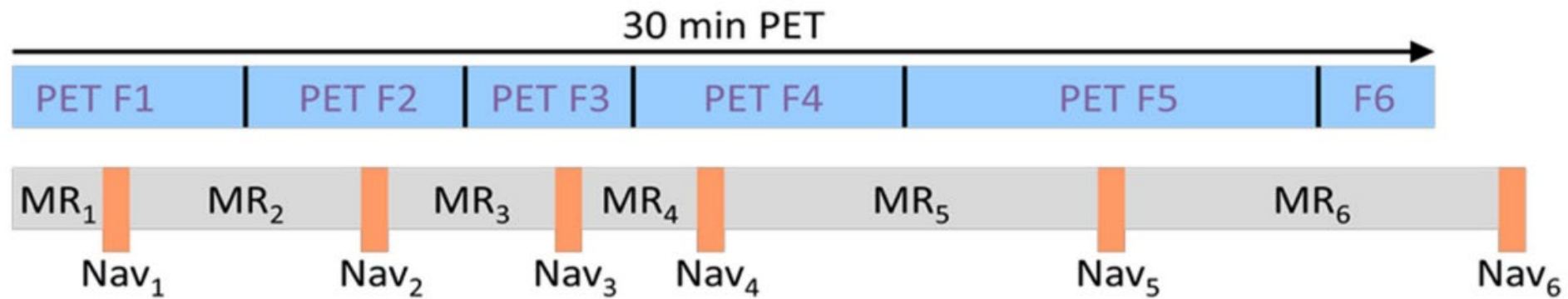
Sagittal

Coronal

Registration

- Intra-scan motion correction usually requires sensors

Part of the acquisition



Wearable sensors



External sensors



Registration

Respiratory and cardiac motion correction for PET/MR

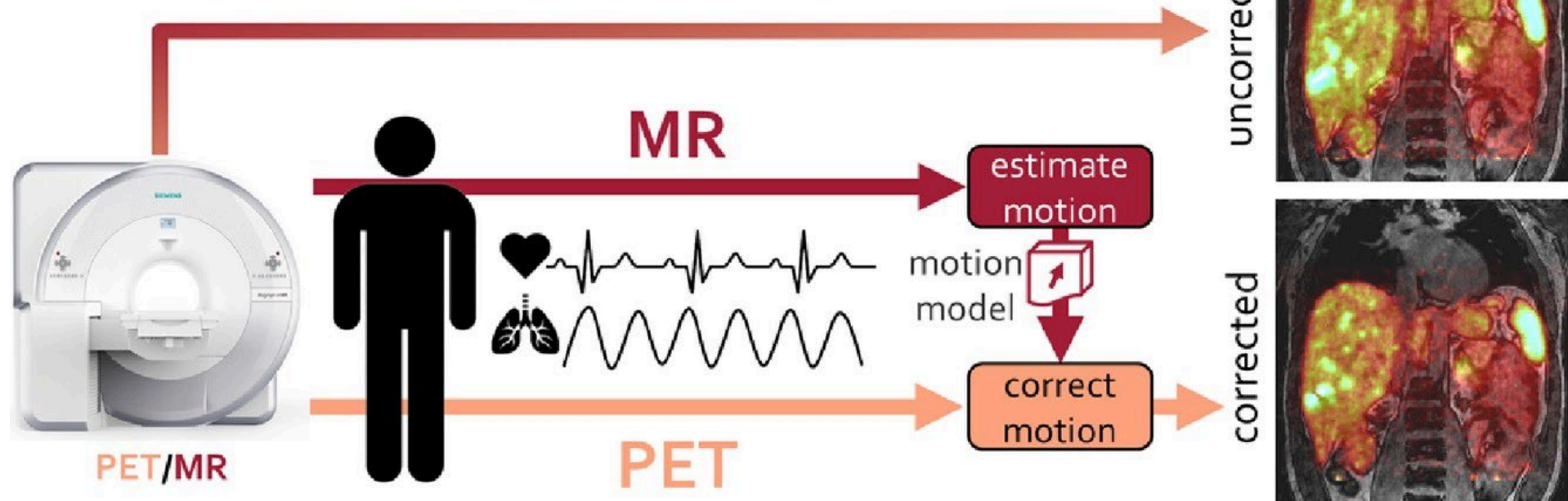


Figure: <https://doi.org/10.1016/j.media.2017.08.002>

Registration

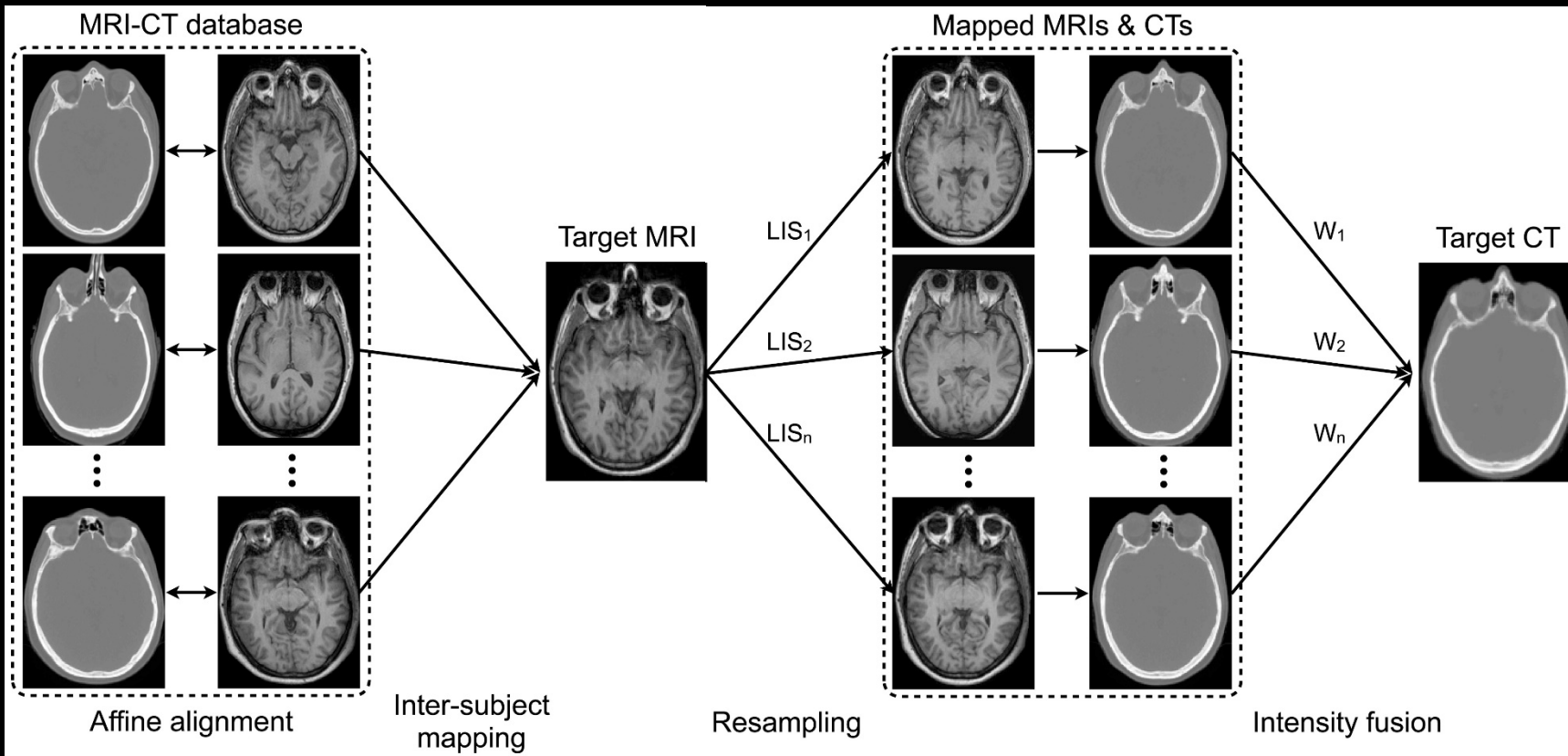
- Goal is to obtain a synthetic CT based on a patient's own MRI

I^{MRI} is target MRI

J_n^{MRI} is warped atlas n

\bar{I} is mean of I

$\sigma(I)$ is standard deviation of I



Simplest solution:

Find best matching warped MRI

$$NCC_n = \frac{1}{N} \frac{\langle I^{MRI} - \bar{I}^{MRI}, J_n^{MRI} - \bar{J}_n^{MRI} \rangle}{\sigma(I^{MRI})\sigma(J_n^{MRI})}$$

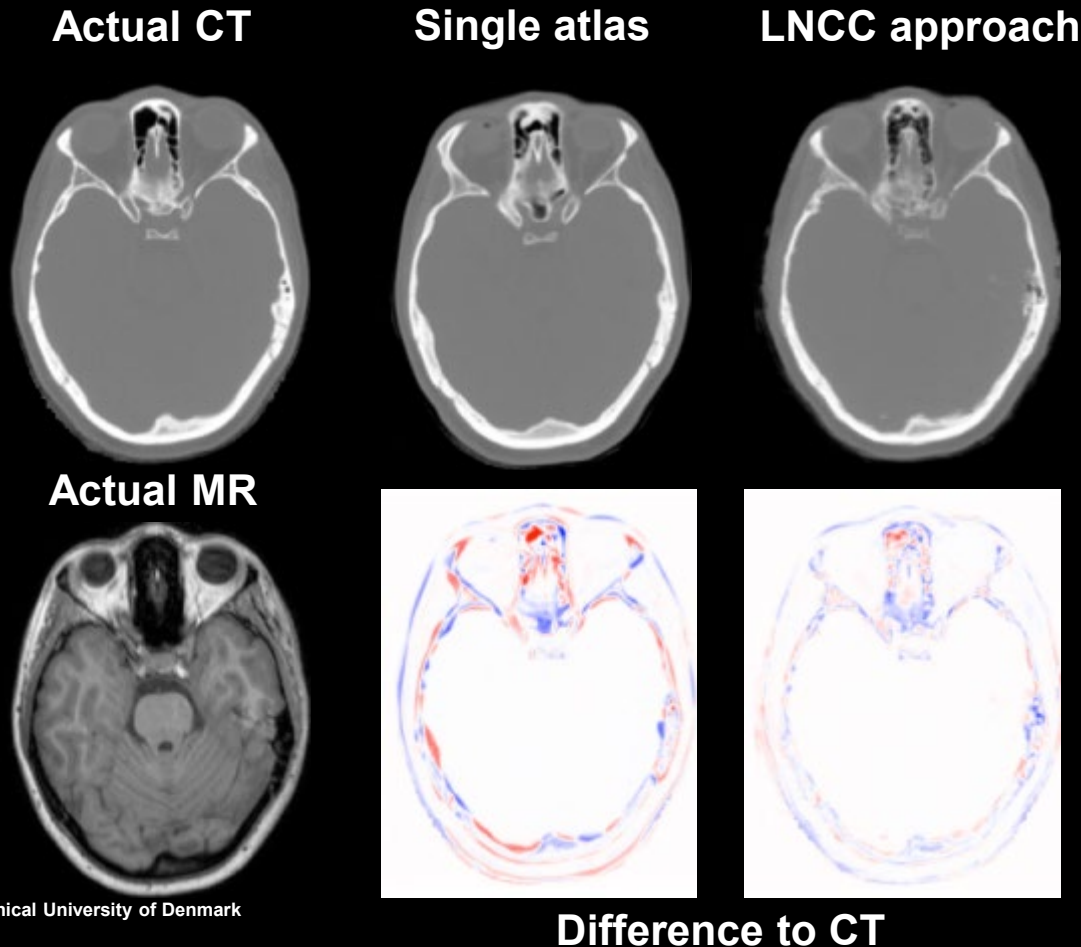
More complex solution:

1. For each voxel, extract patch and compute local NCC (LNCC)
2. Rank the patches based on their LNCC
3. Fuse the CT values based on their ranks (higher rank = higher weight)

Registration

I^{MRI} is target MRI
 J_n^{MRI} is warped atlas n
 \bar{I} is mean of I
 $\sigma(I)$ is standard deviation of I

- Goal is to obtain a synthetic CT based on a patient's own MRI



Simplest solution:
 Find best matching warped MRI

$$NCC_n = \frac{1}{N} \frac{\langle I^{MRI} - \bar{I}^{MRI}, J_n^{MRI} - \bar{J}_n^{MRI} \rangle}{\sigma(I^{MRI})\sigma(J_n^{MRI})}$$

More complex solution:

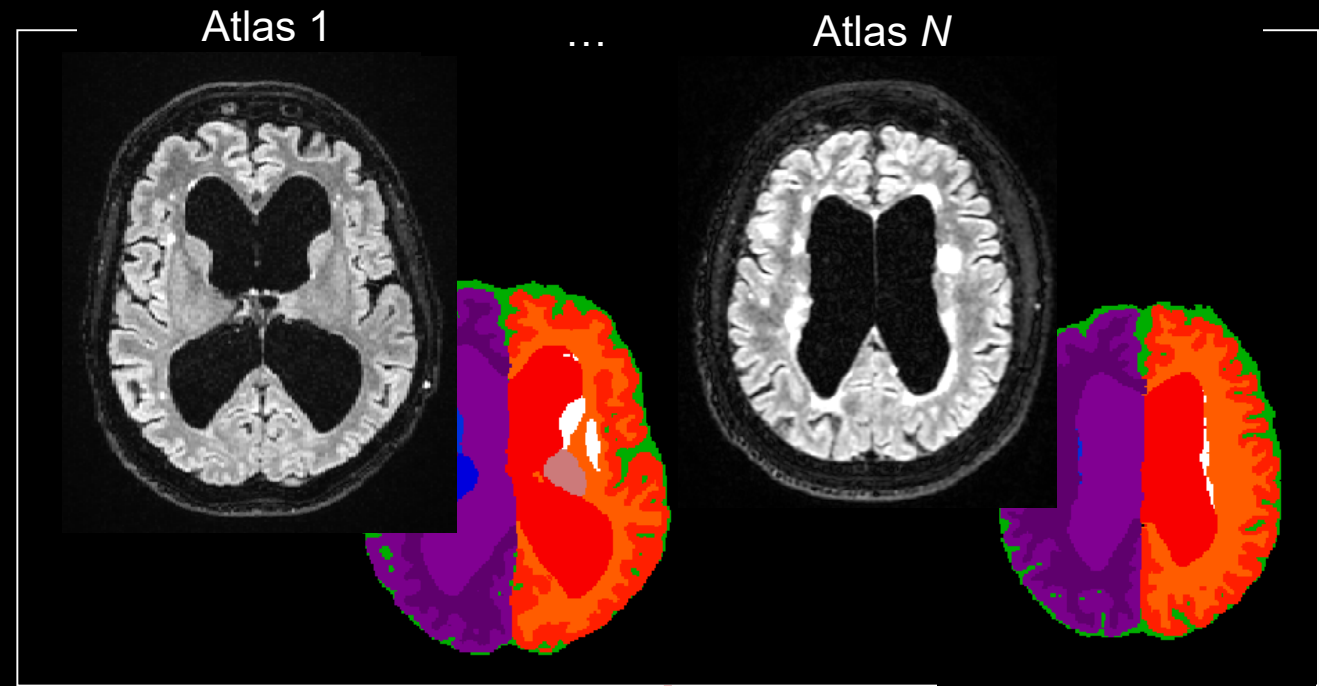
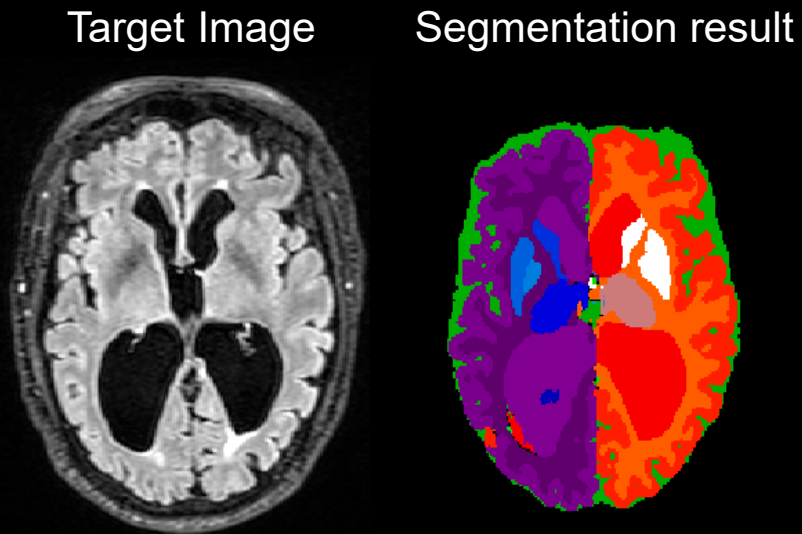
1. For each voxel, extract patch and compute local NCC (LNCC)
2. Rank the patches based on their LNCC
3. Fuse the CT values based on their ranks (higher rank = higher weight)

Detection

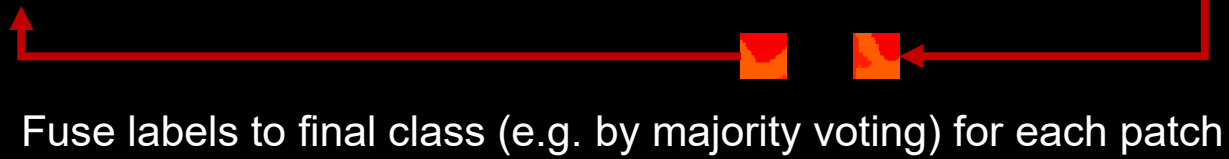
- Segmentation
- Detection
- Tracking

Segmentation

- Label fusion



Warp each atlas to target image

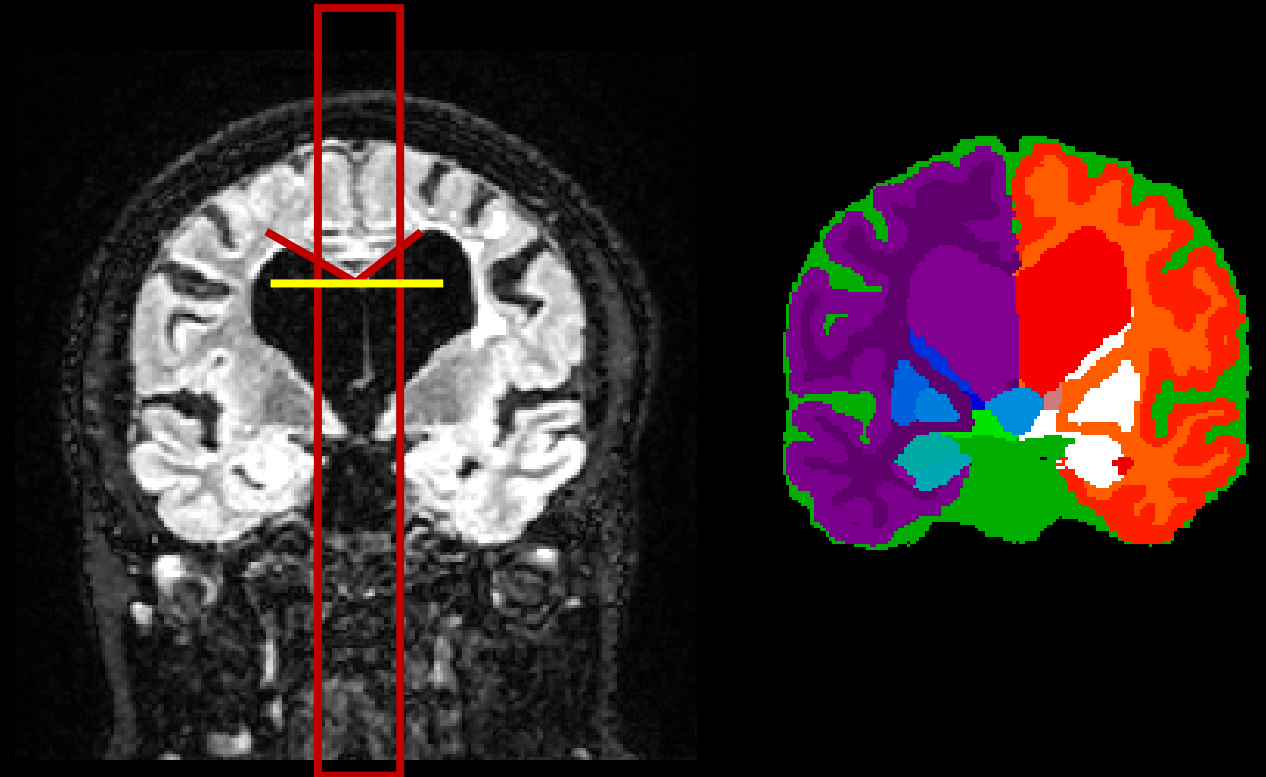


Quiz 4

- The 10 estimates for a class label are found after registration.
 - [1, 5, 2, 1, 2, 5, 4, 5, 2, 2]
- Using majority voting, what is the final predicted class?
 - Answers:
 - 1
 - 2
 - 4
 - 5

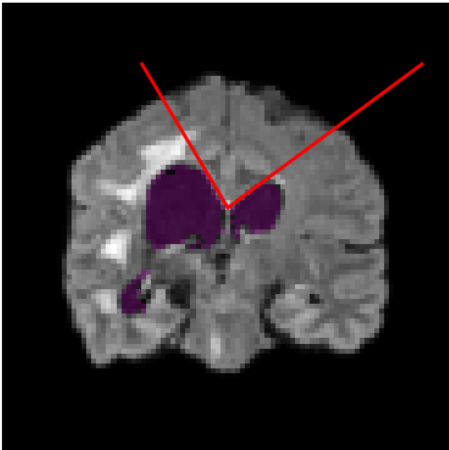
Detection

- Determine the Callosal angle
- Steps
 1. Align MRI to standard space to select standard center slice
 2. Determine first row without brain tissue in center columns
 3. Fit a line to brain tissue points for each side
 4. Determine angle between lines

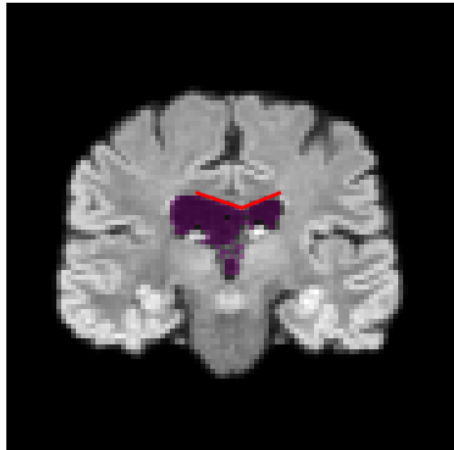


Detection

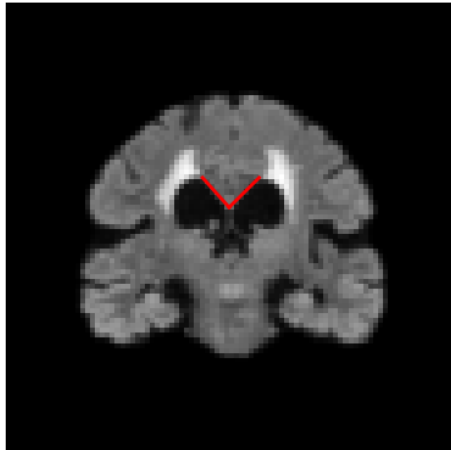
Callosal Angle: 84



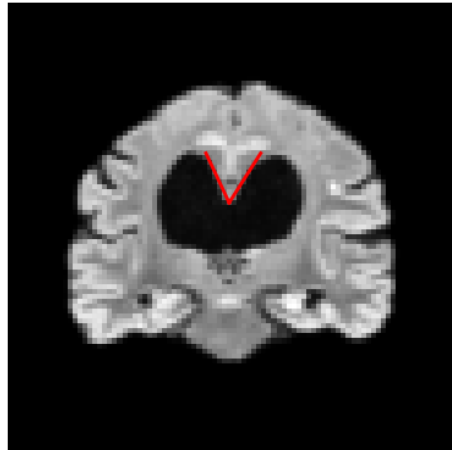
Callosal Angle: 140



Callosal Angle: 87



Callosal Angle: 57



Callosal Angle: 66

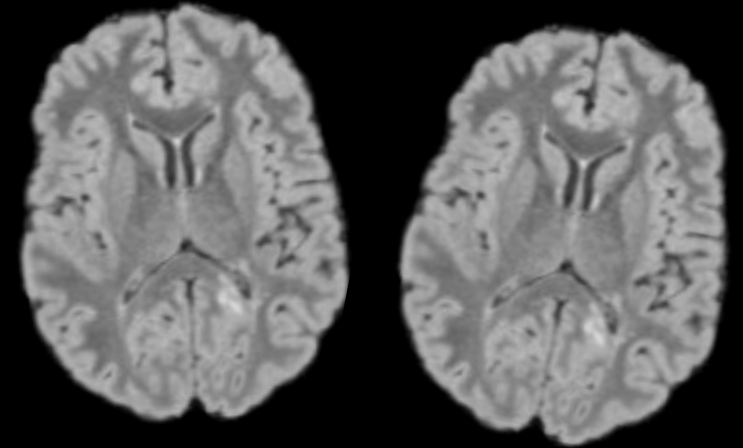


Tracking

- Tracking of objects over time to detect progression

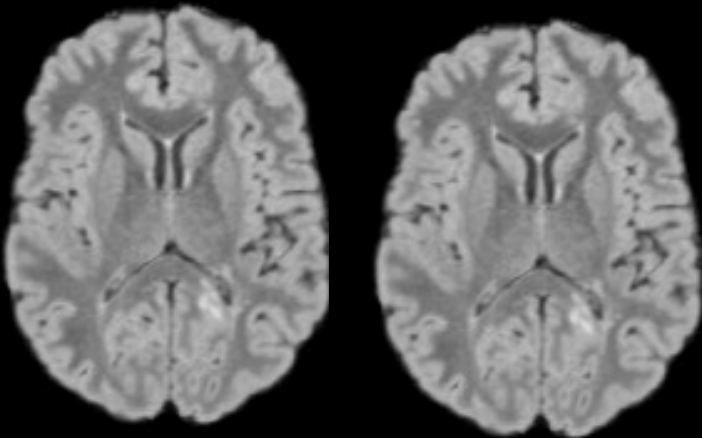
Baseline

Follow-up



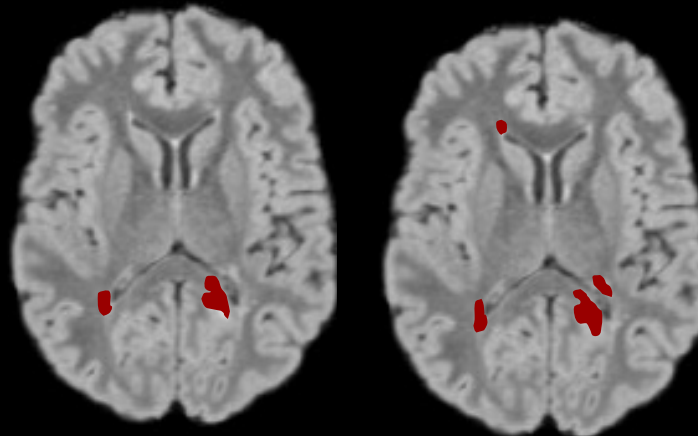
Step 1

Register images



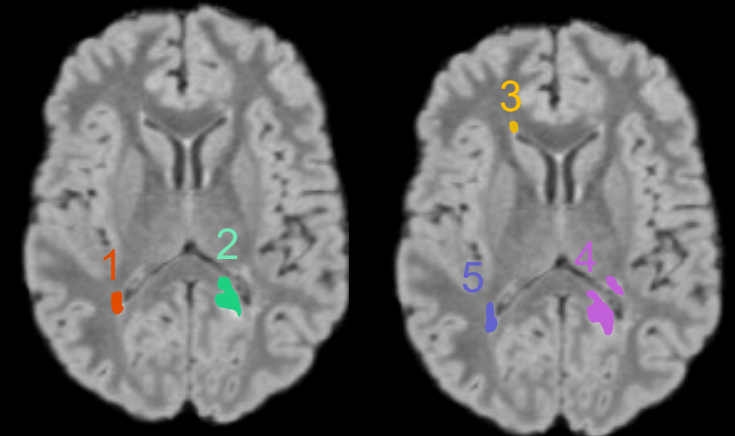
Step 2

Segment lesions



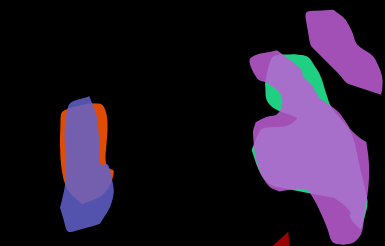
Step 3

Connected component analysis



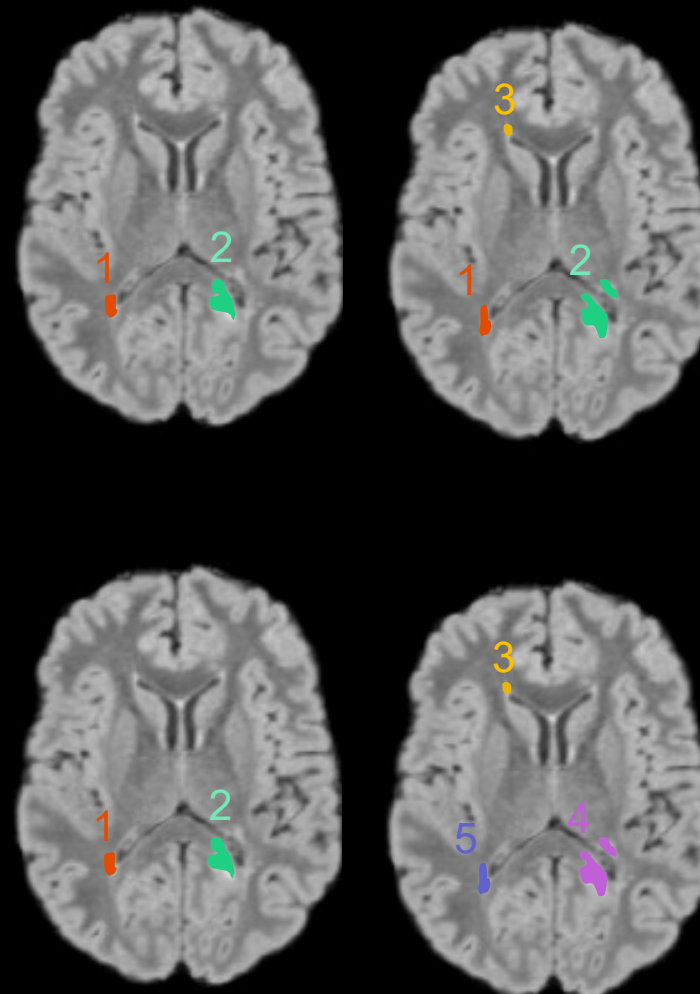
Tracking

New cluster



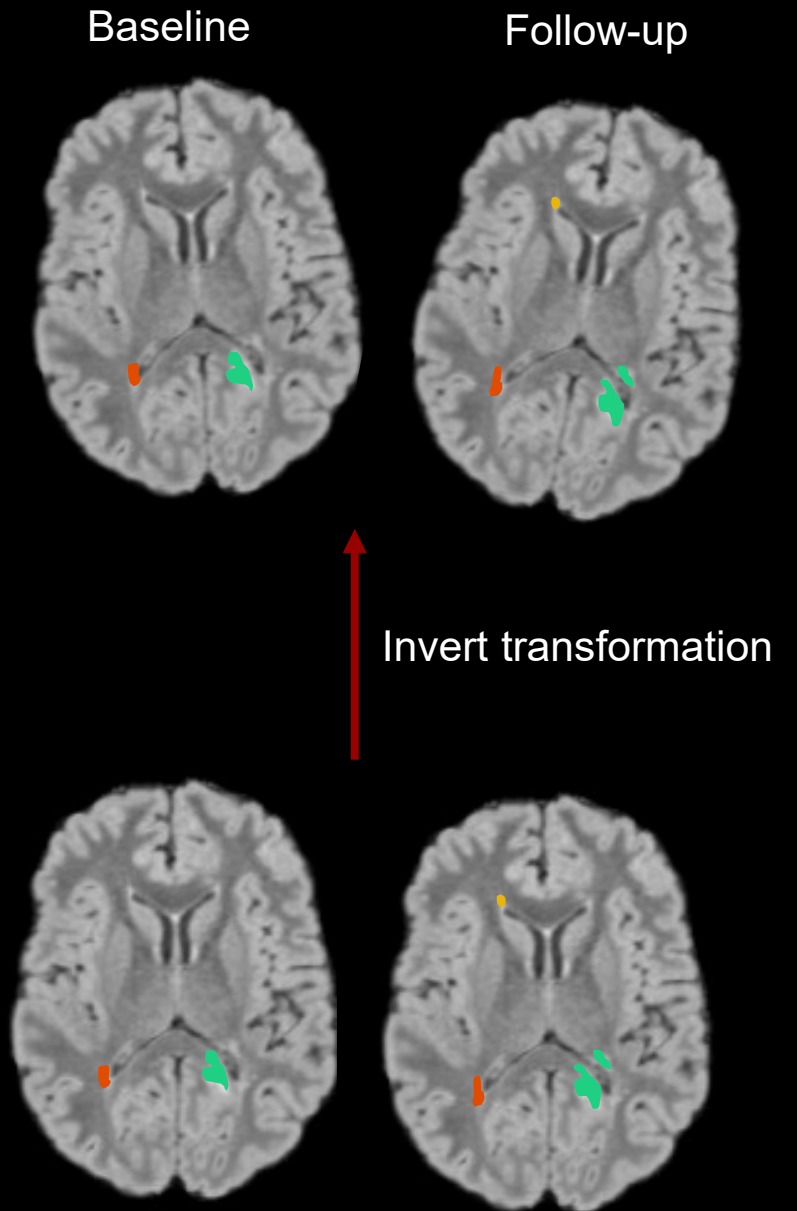
Overlapping clusters

Step 4
Global remapping



Tracking

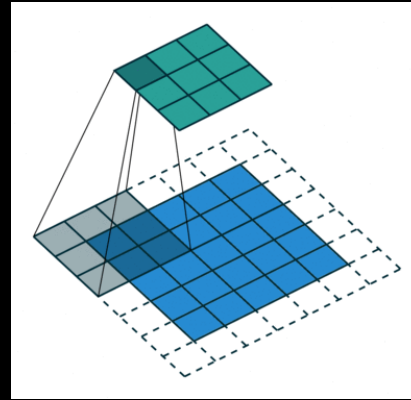
- Tracking of objects over time to detect progression



Classification (and more)

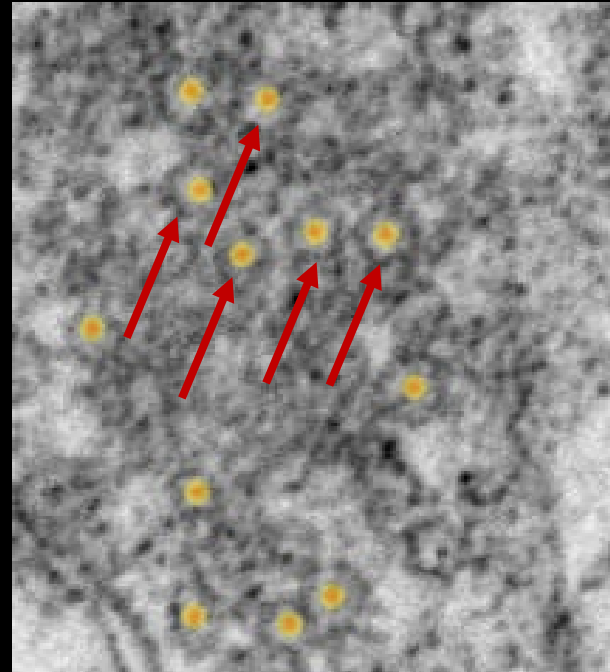
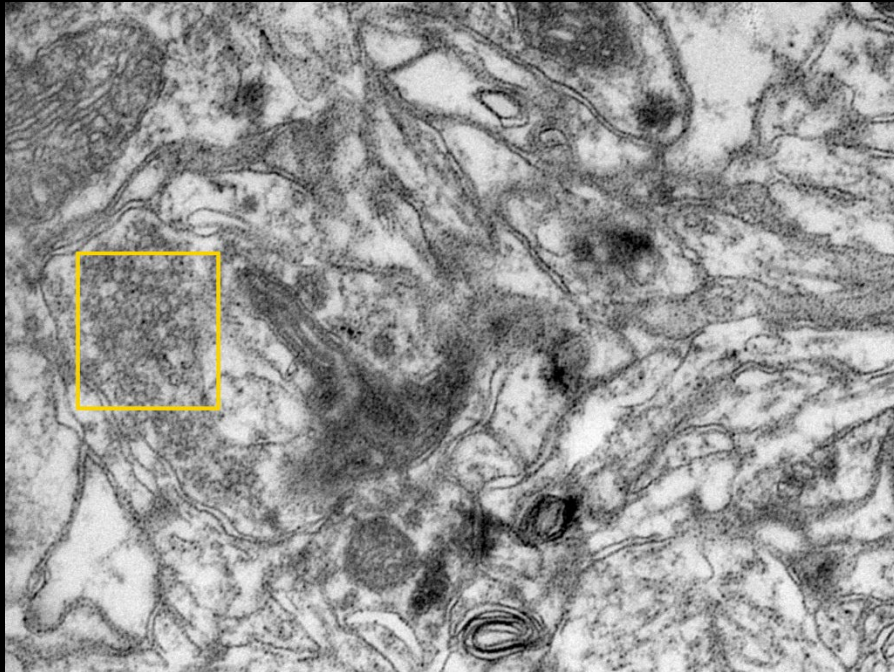
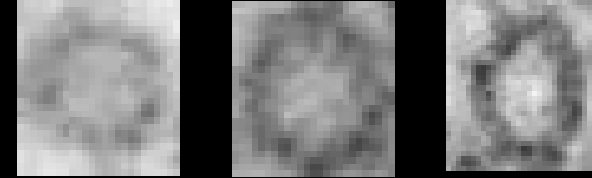
- Template matching
- Feature engineering
- Random Forest
- Active Shape Models
- Active Contours

Template matching

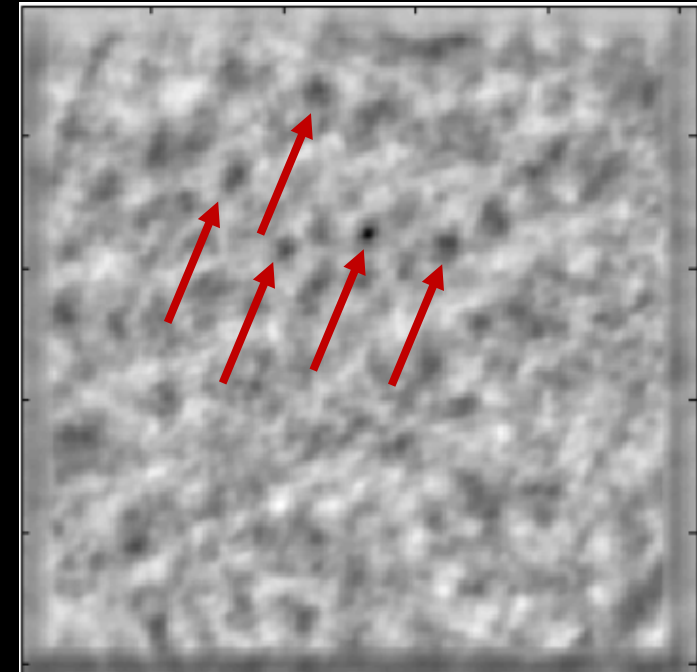


$$g(x, y) = \sum_{j=-R}^R \sum_{i=-R}^R h(i, j) \cdot f(x + i, y + j)$$

Examples of h :



Reference



Resulting g

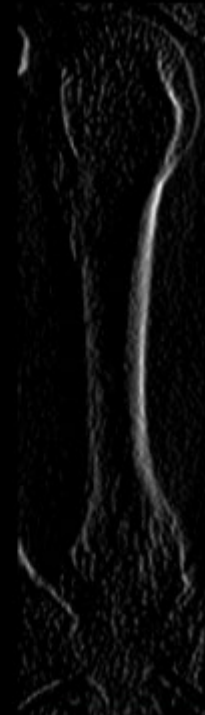
Feature engineering

What is relevant to know about this image to classify each voxel/pixel?

- Edges?
- Shapes?



Original



Prewitt



Circle like



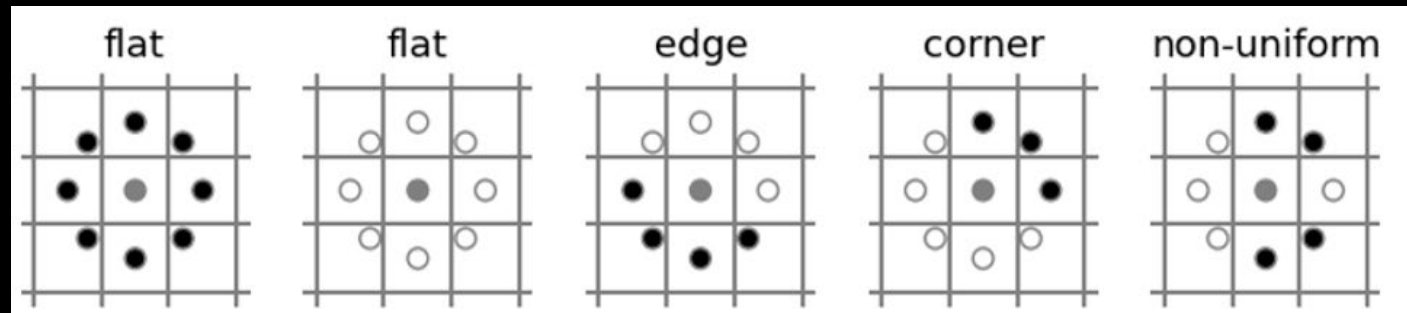
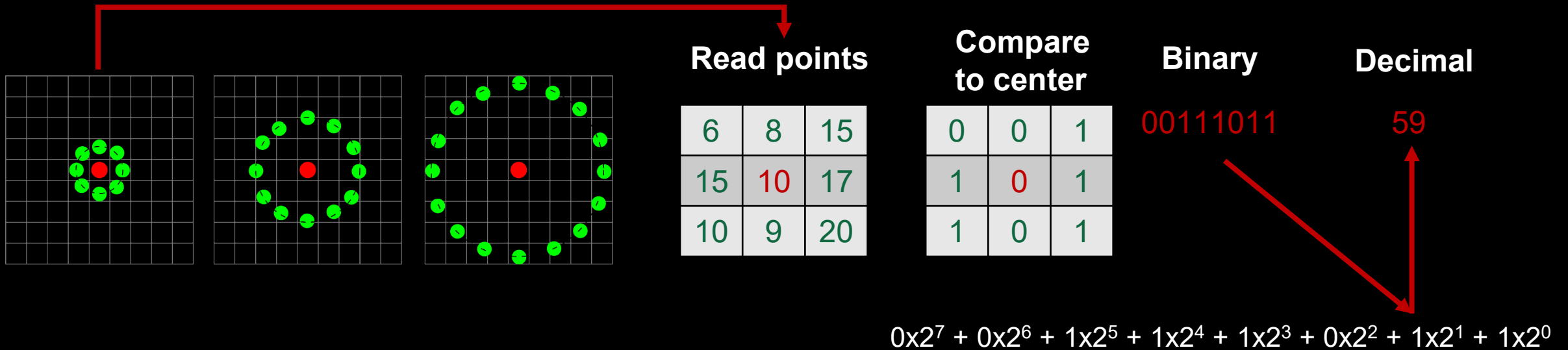
Not circle like

Week #4, Filtering

Week #5, Blob features

Feature engineering – Local Binary Patterns

Tunable parameters include radius (distance between center and points) and number of points on grid



Quiz 5

- Given the read matrix, what is the calculated LBP value
 - 163
 - 167
 - 171
 - 180

Read matrix

6	4	6
15	5	4
10	9	3

Quiz 5

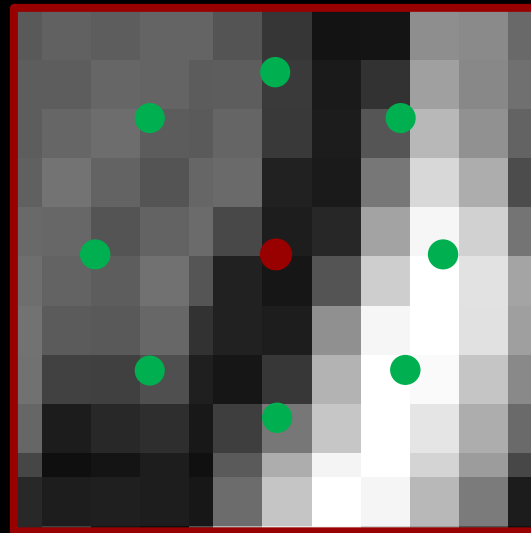
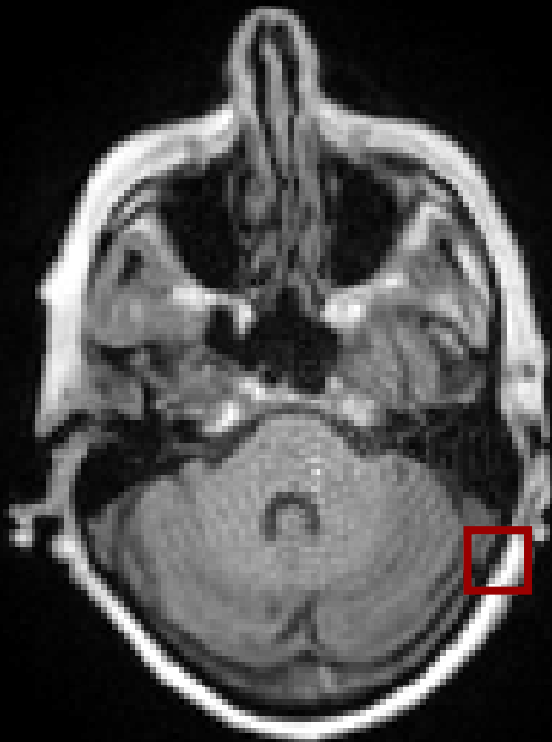
- Given the read matrix, what is the calculated LBP value
 - 163
 - 167
 - 171
 - 180

Read matrix

6	4	6
15	5	4
10	9	3

$$10100111 = 128 + 32 + 4 + 2 + 1 = 167$$

Feature engineering – Local Binary Patterns



Read points

30	20	35
28	10	41
15	37	45

Compare to center

1	1	1
1	0	1
1	1	1

Binary

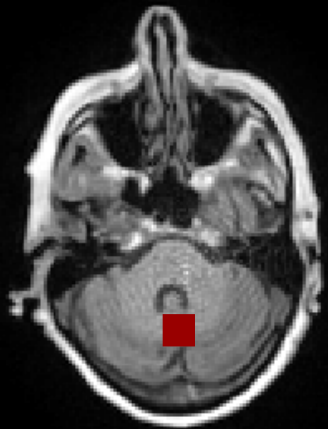
11111111

Decimal

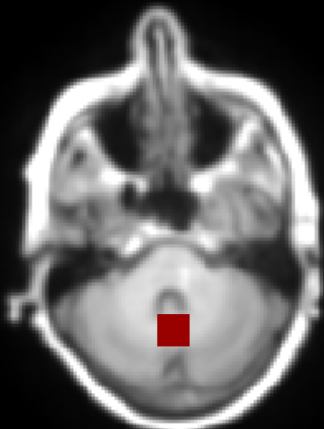
255



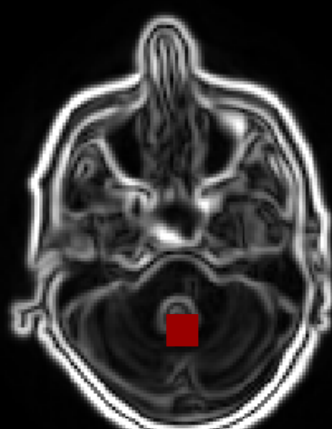
Feature engineering



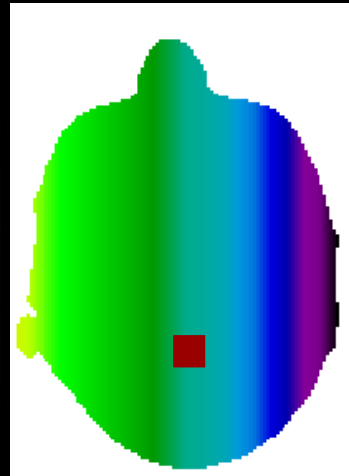
Normalized



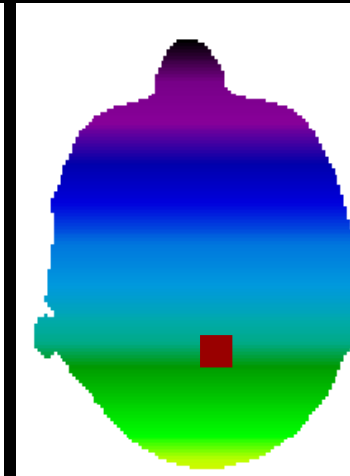
Blurred



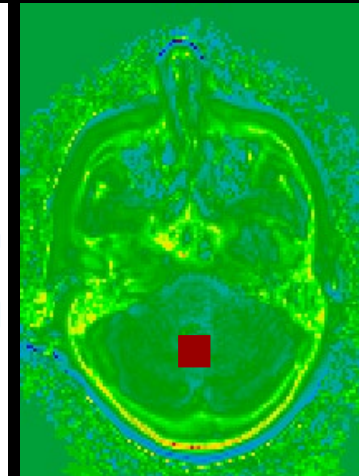
Gradient
magnitude



Spatial-x



Spatial-y

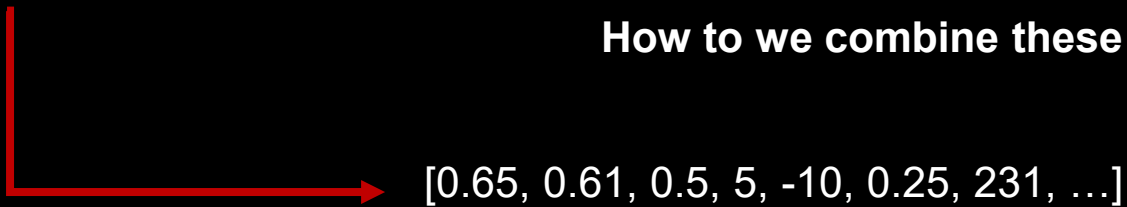


R2*

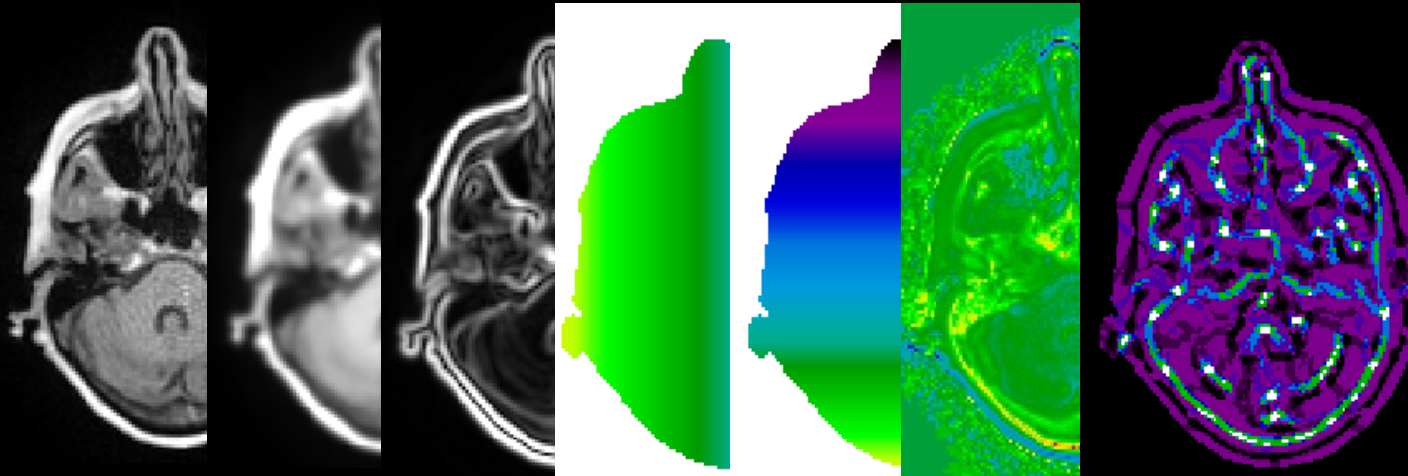


LBP

How to we combine these into a voxel classification model?



Feature engineering



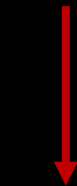
Repeat for all voxels:

[0.65, 0.61, 0.5, 5, -10, 0.25, 231, ...]
 [0.45, 0.66, 0.4, 6, -12, 0.24, 251, ...]
 ...
 ...
 [0.87, 0.41, 0.1, 2, 25, 0.55, 131, ...]

Normalize to 0-1 range

[0.61, 0.59, 0.63, 0.5, 0.11, 0.25, 0.88, ...]
 [0.41, 0.65, 0.45, 0.6, 0.08, 0.24, 0.95, ...]
 ...
 ...
 [0.81, 0.38, 0.12, 0.2, 0.31, 0.55, 0.45, ...]

Reference value
for each voxel

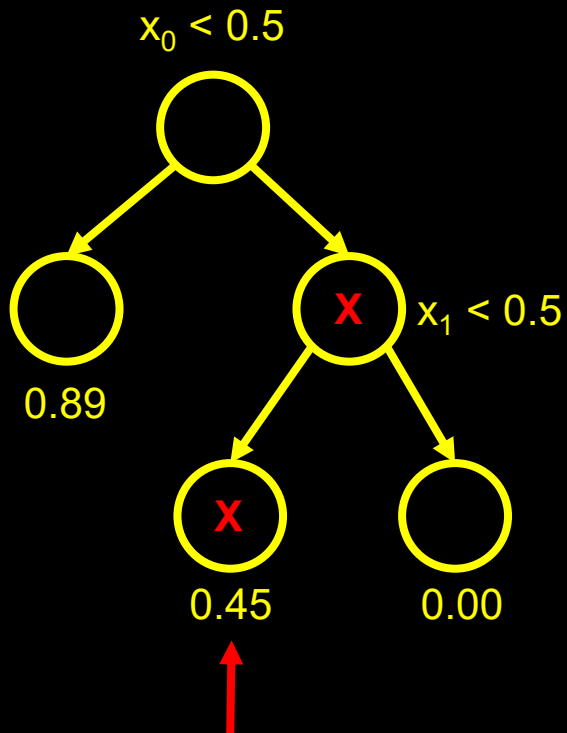


[0.00]
 [0.89]
 ...
 [0.45]

[id]	[x ₀ , x ₁ , ...]	[y]
[0]	[0.61, 0.59, 0.63, 0.5, 0.11, 0.25, 0.88, ...]	[0.00]
[1]	[0.41, 0.65, 0.45, 0.6, 0.08, 0.24, 0.95, ...]	[0.89]
...
[n]	[0.81, 0.38, 0.12, 0.2, 0.31, 0.55, 0.45, ...]	[0.45]

Random Forest

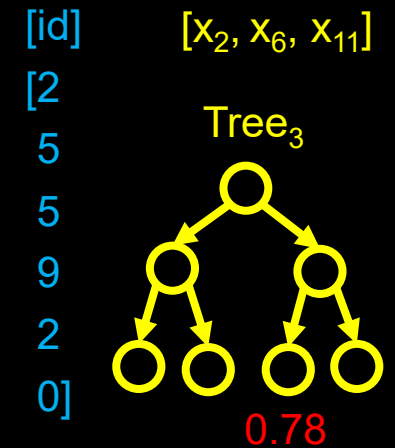
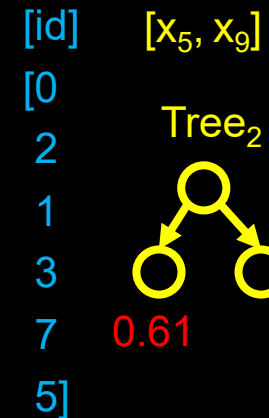
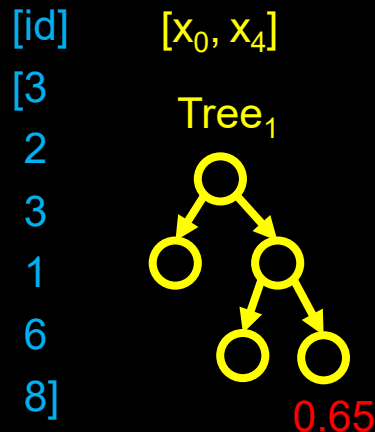
Decision tree



- Multiple trees make a forest
- Why random?
 - Each tree sees a random data sample with replacement (Bootstrap)
 - Each tree sees a random subset of the variables

New datapoint: **[0.65, 0.33, ...,]**

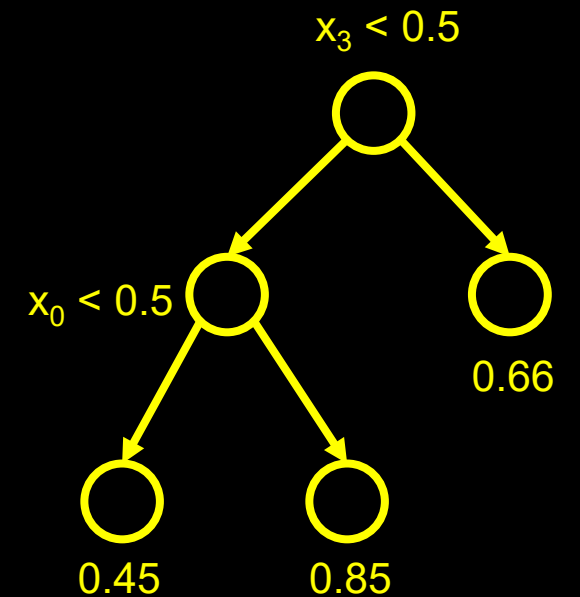
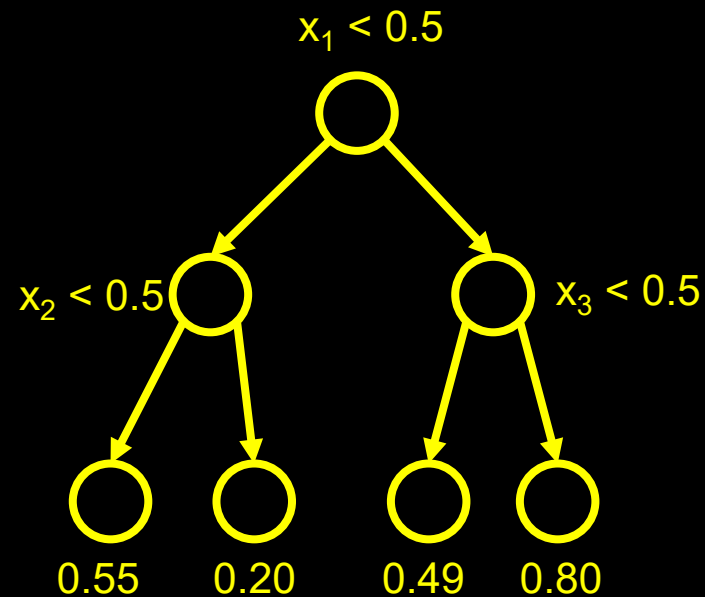
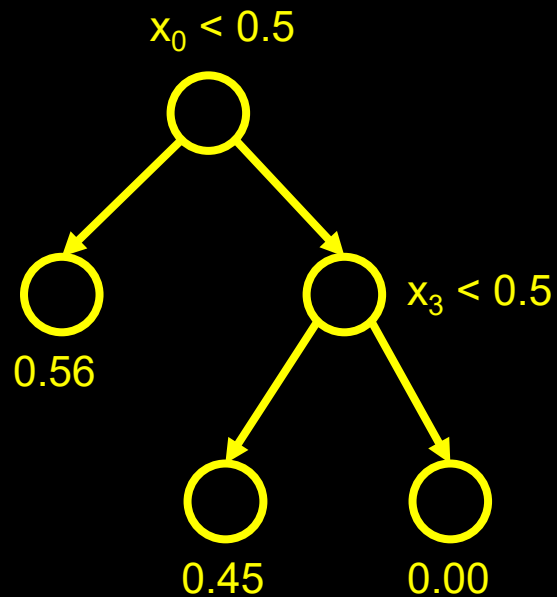
Mean of results (Aggregating): $\bar{y} = \frac{1}{n} \sum_i \bar{y}_i = \frac{1}{3} (0.65 + 0.61 + 0.78) = \underline{0.68}$



Quiz 6

- Using the 3 trained trees below, what is the predicted value after aggregating the output?
- Input data: [0.49, 0.56, 0.99, 0.32]
- Options:

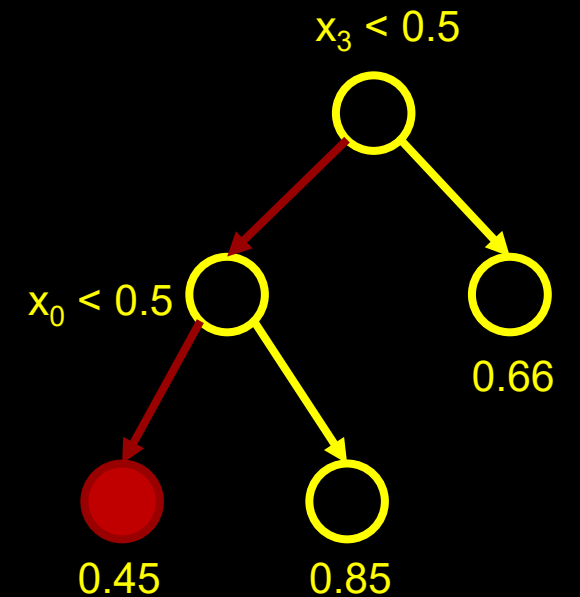
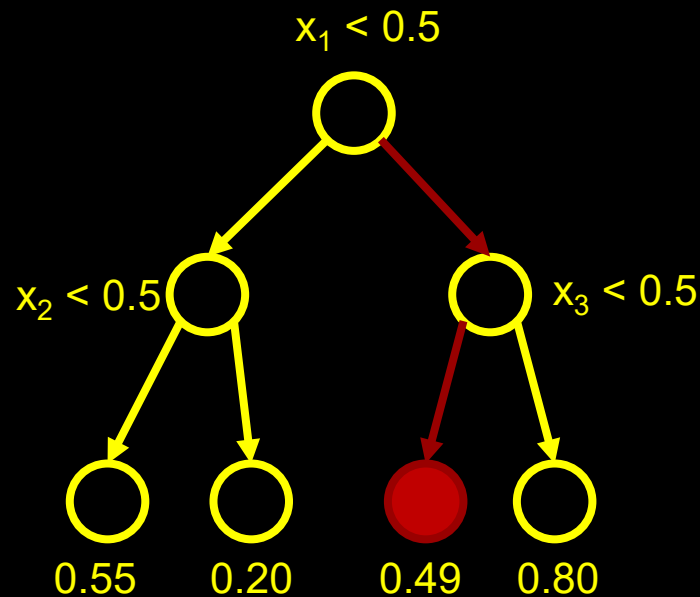
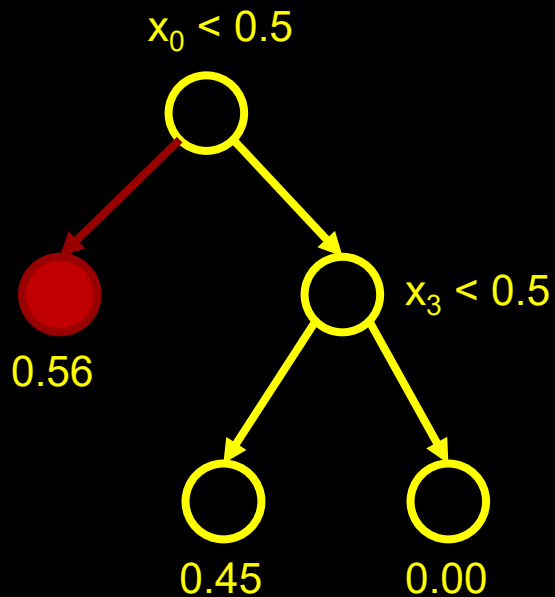
- 0.45
- 0.50
- 0.80
- 0.48



Quiz 6

- Using the 3 trained trees below, what is the predicted value after aggregating the output?
- Input data: [0.49, 0.56, 0.99, 0.32]
- Options:

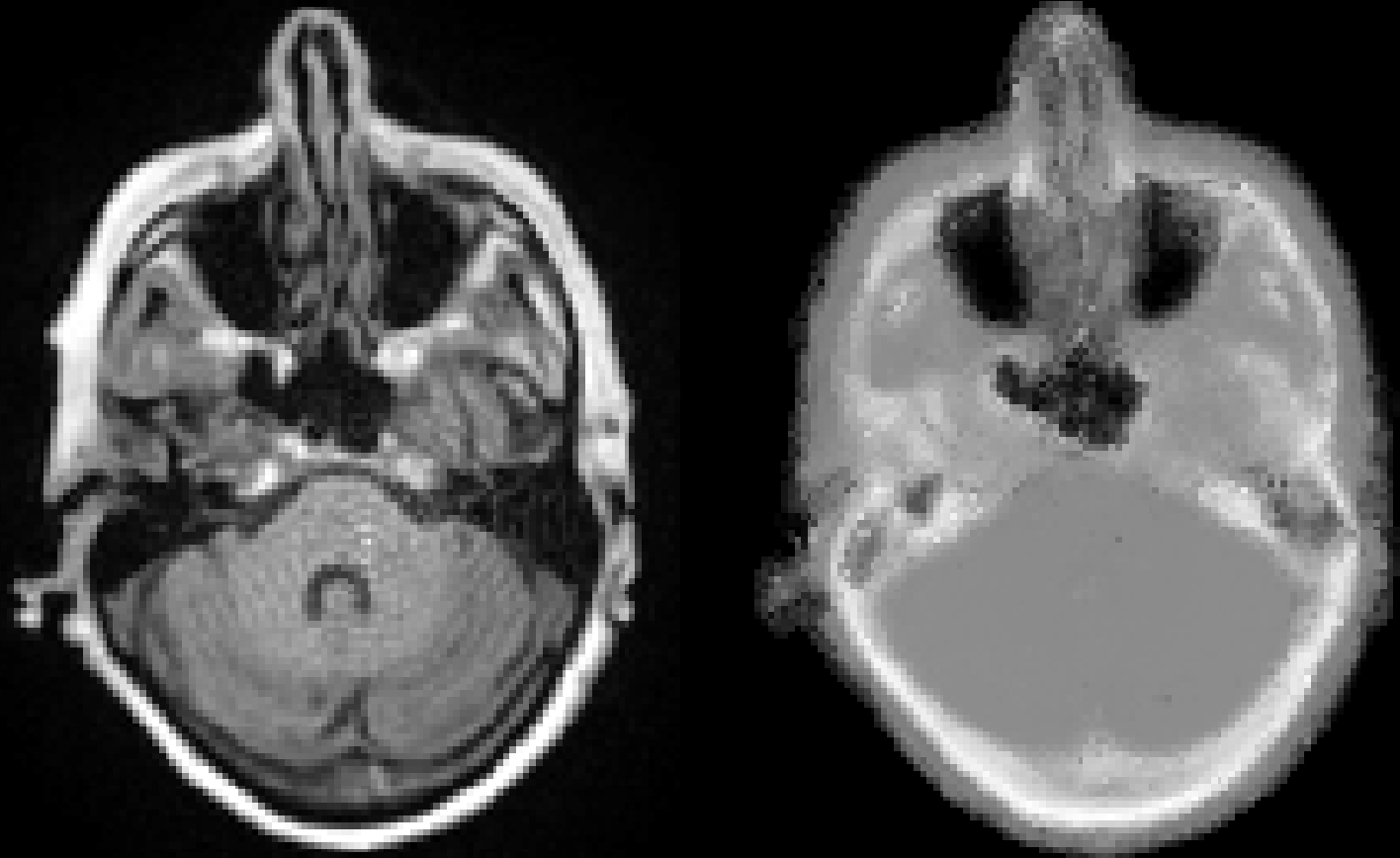
- 0.45
- 0.50
- 0.80
- 0.48



$$(0.56 + 0.49 + 0.45)/3 = 0.5$$

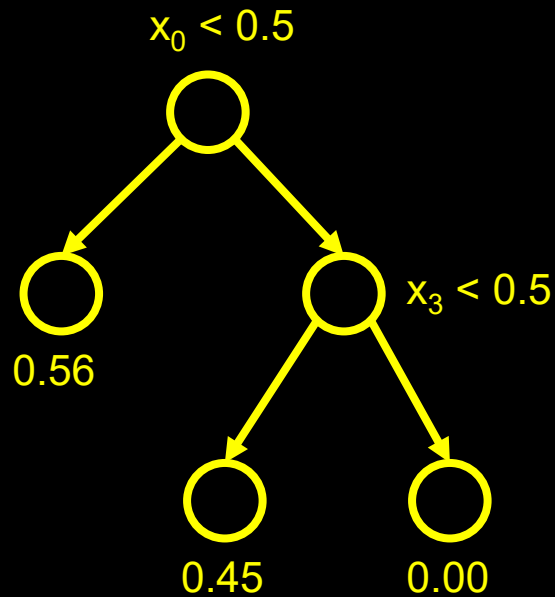
Random forest

- Example output:
 - 100 trees
 - n=25 patients
 - Features from
 - Original and filtered images
 - Edge enhanced
 - R2*
 - LBP
- Trained with RandomForestRegressor from sklearn



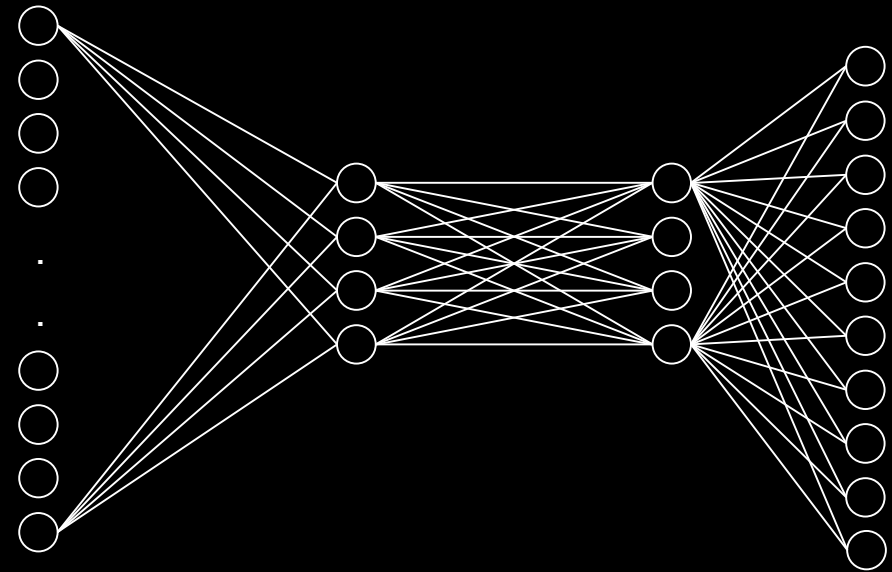
Increasing complexity..

Random Forest



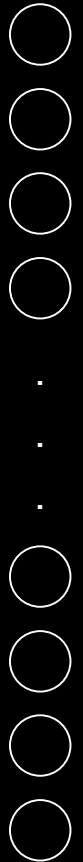
(Potential for) high level of interpretability

Neural network



Limited interpretability

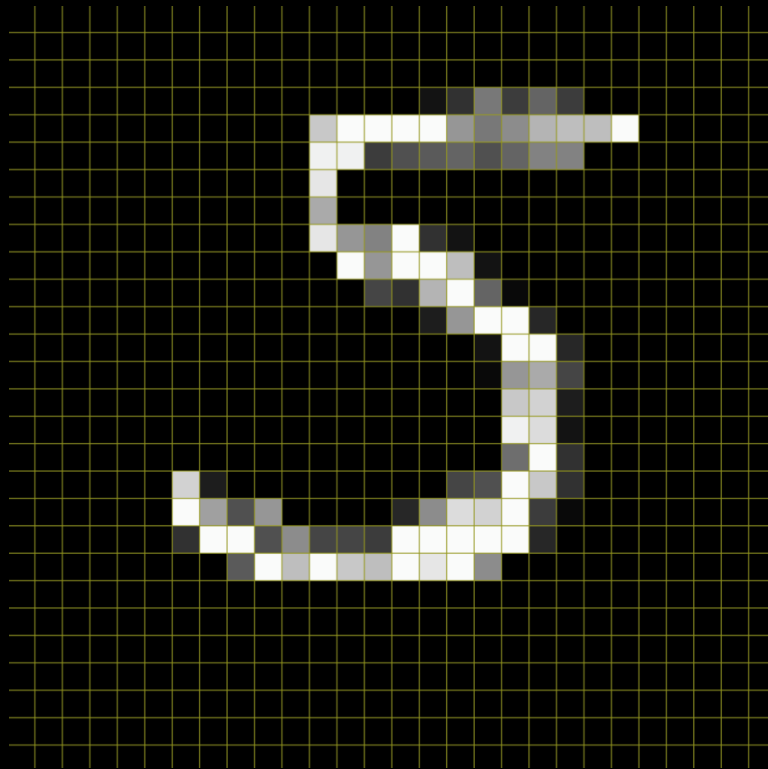
Neural Networks



Vectorize

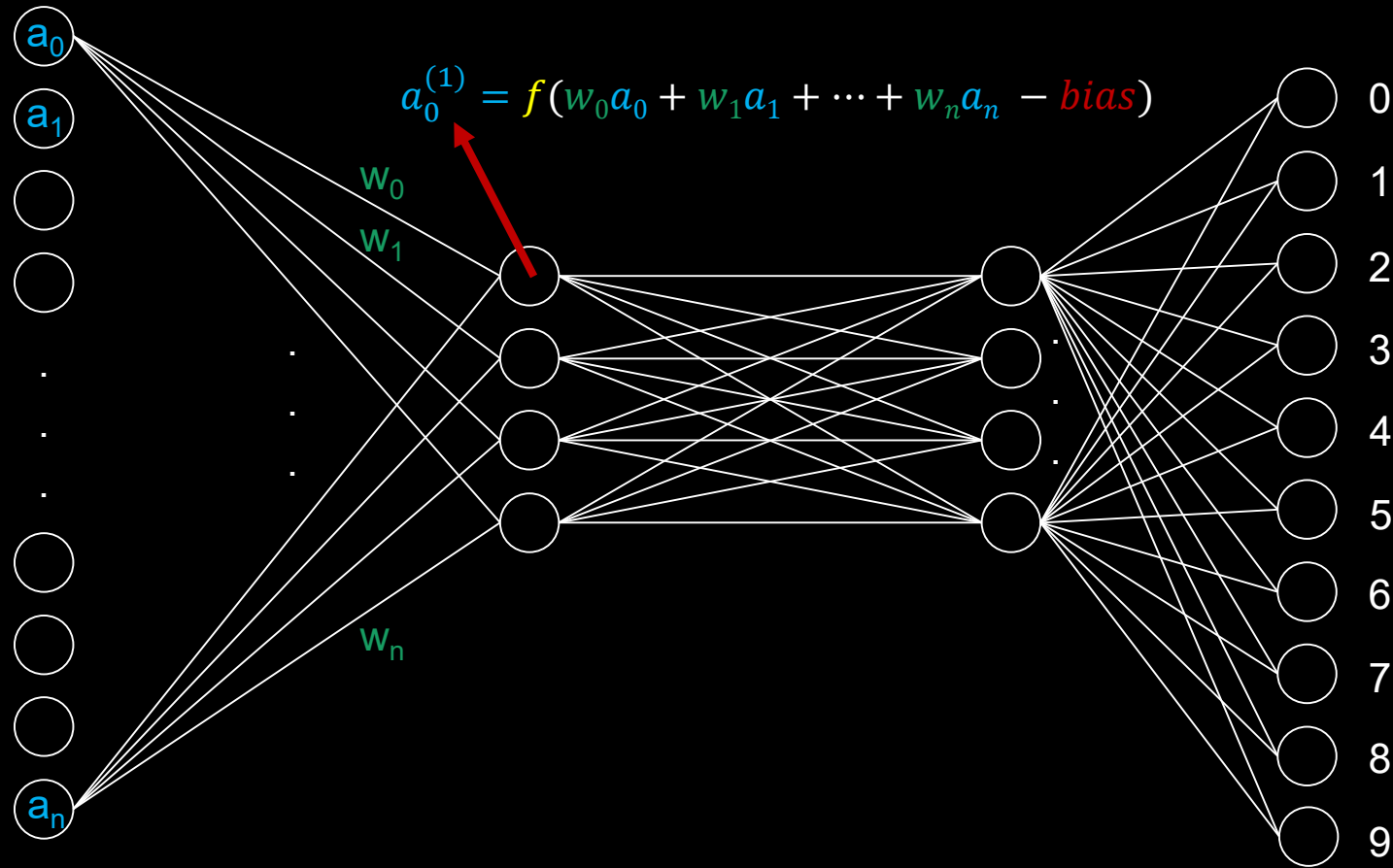


$28 \times 28 = 784$



- 0
- 1
- 2
- 3
- 4
- 5
- 6
- 7
- 8
- 9

Neural Networks



- Each neuron contains a value, its "activation"
 - The values in the input are the pixel values
 - The value at the last output layer represents the likelihood of that digit
 - f is an activation function (e.g. sigmoid)

weights: $784 \times 4 + 4 \times 4 + 4 \times 10$

biases: $4 + 4 + 10$

Total parameters: 3,210

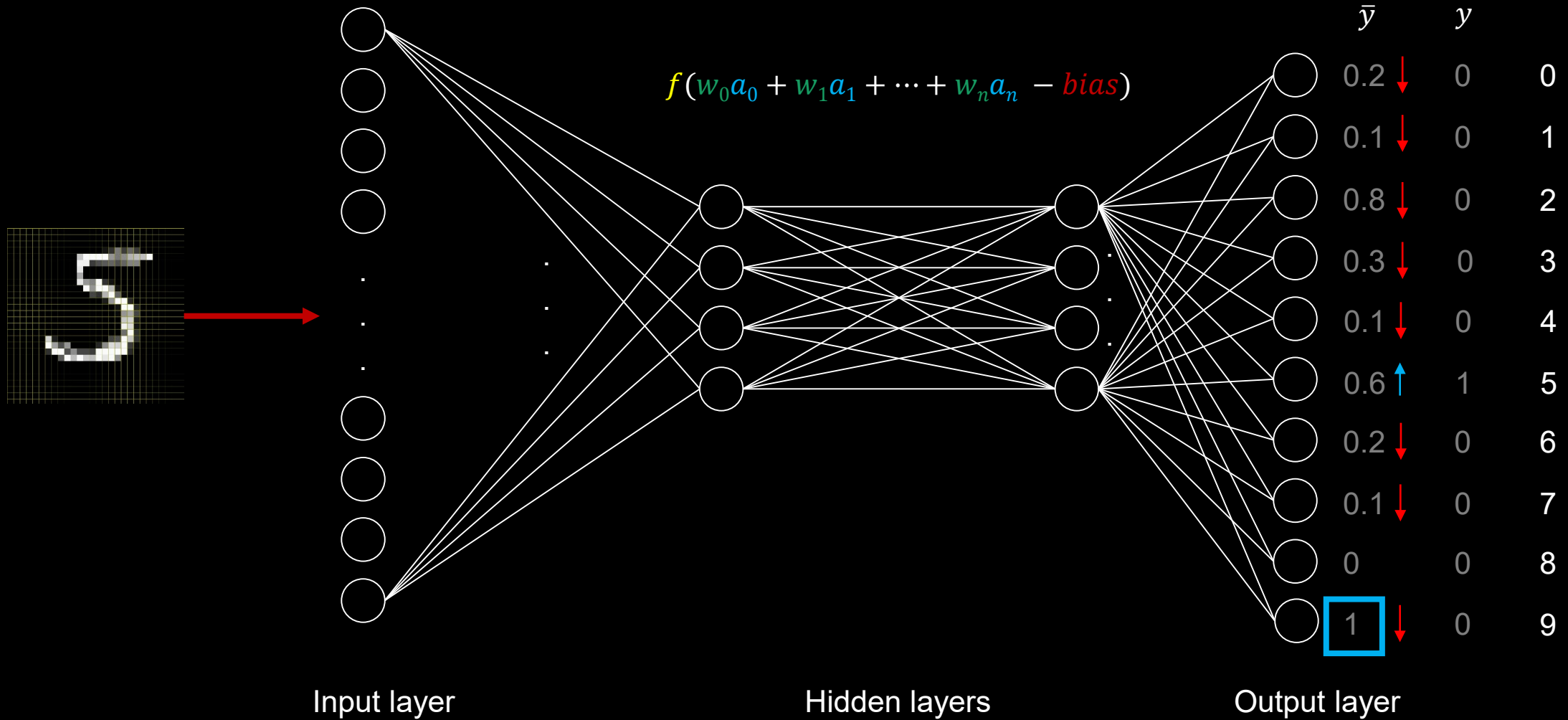
Input layer

Hidden layers

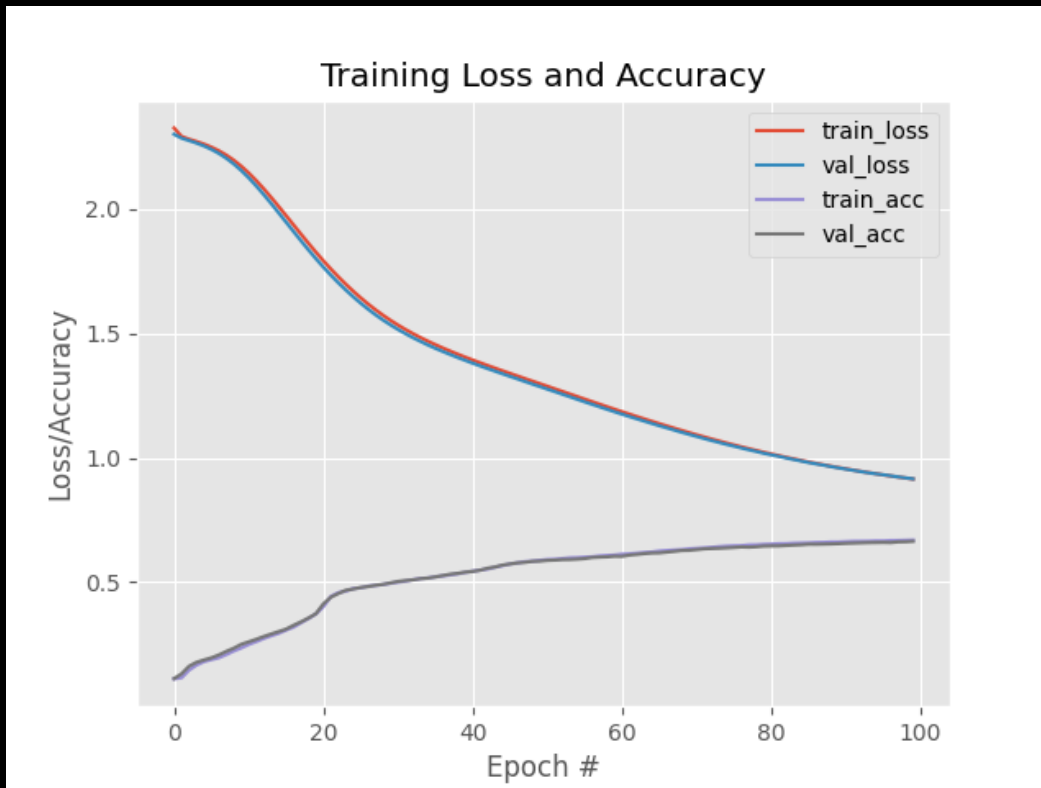
Output layer

$$\sum (\bar{y} - y)^2$$

Neural Networks



Neural Networks



Load and prepare data

```
from tensorflow.keras.datasets import mnist
((trainX, trainY), (testX, testY)) = mnist.load_data()

# Vectorize
trainX = trainX.reshape((trainX.shape[0], 28 * 28 * 1))
testX = testX.reshape((testX.shape[0], 28 * 28 * 1))
# scale data to the range of [0, 1]
trainX = trainX.astype("float32") / 255.0
testX = testX.astype("float32") / 255.0
```

Define model

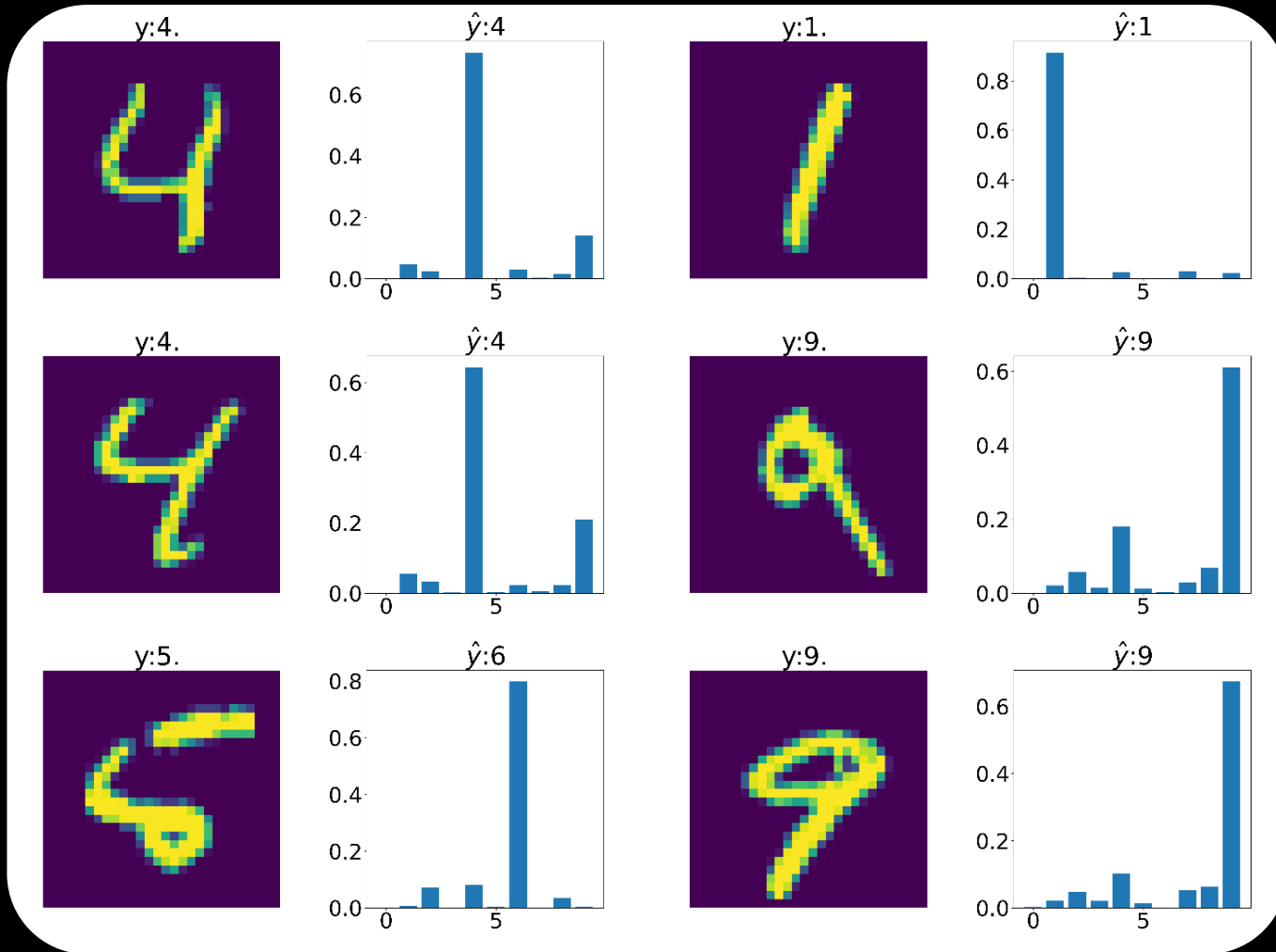
```
model = Sequential()
model.add(Dense(4, input_shape=(784,), activation="sigmoid"))
model.add(Dense(4, activation="sigmoid"))
model.add(Dense(10, activation="softmax"))
```

Train model

```
model.compile(
    loss="categorical_crossentropy",
    optimizer=SGD(0.01),
    metrics=["accuracy"])

model.fit(
    trainX, trainY, validation_data=(testX, testY),
    epochs=100, batch_size=128)
```

Neural Networks



Load and prepare data

```
from tensorflow.keras.datasets import mnist
((trainX, trainY), (testX, testY)) = mnist.load_data()

# Vectorize
trainX = trainX.reshape((trainX.shape[0], 28 * 28 * 1))
testX = testX.reshape((testX.shape[0], 28 * 28 * 1))

# scale data to the range of [0, 1]
trainX = trainX.astype("float32") / 255.0
testX = testX.astype("float32") / 255.0
```

Define model

```
model = Sequential()
model.add(Dense(4, input_shape=(784,), activation="sigmoid"))
model.add(Dense(4, activation="sigmoid"))
model.add(Dense(10, activation="softmax"))
```

Train model

```
model.compile(
    loss="categorical_crossentropy",
    optimizer=SGD(0.01),
    metrics=["accuracy"])

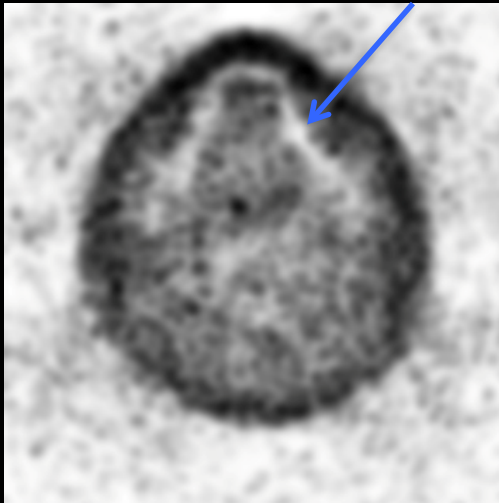
model.fit(
    trainX, trainY, validation_data=(testX, testY),
    epochs=100, batch_size=128)
```

Active Shape Models – and more

Motivation: Artifacts in umaps result in loss of quantitative accuracy



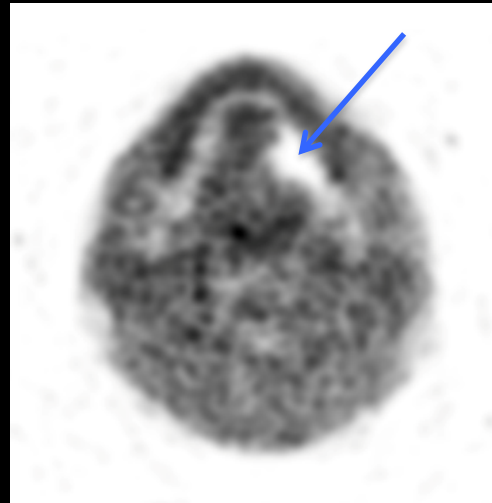
μ -map



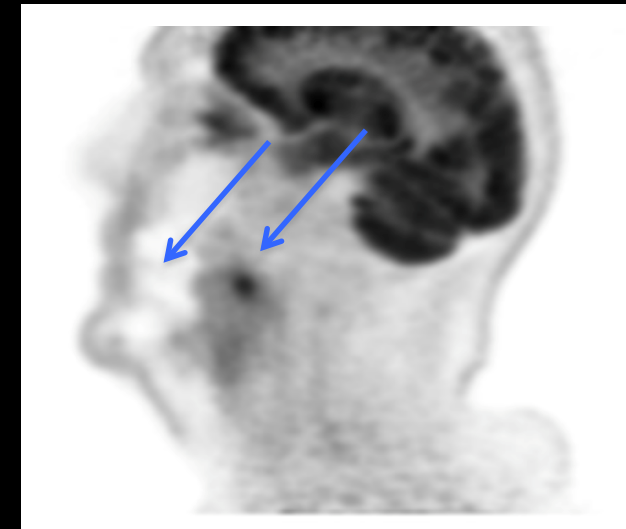
NAC-PET_{MR}



μ -map

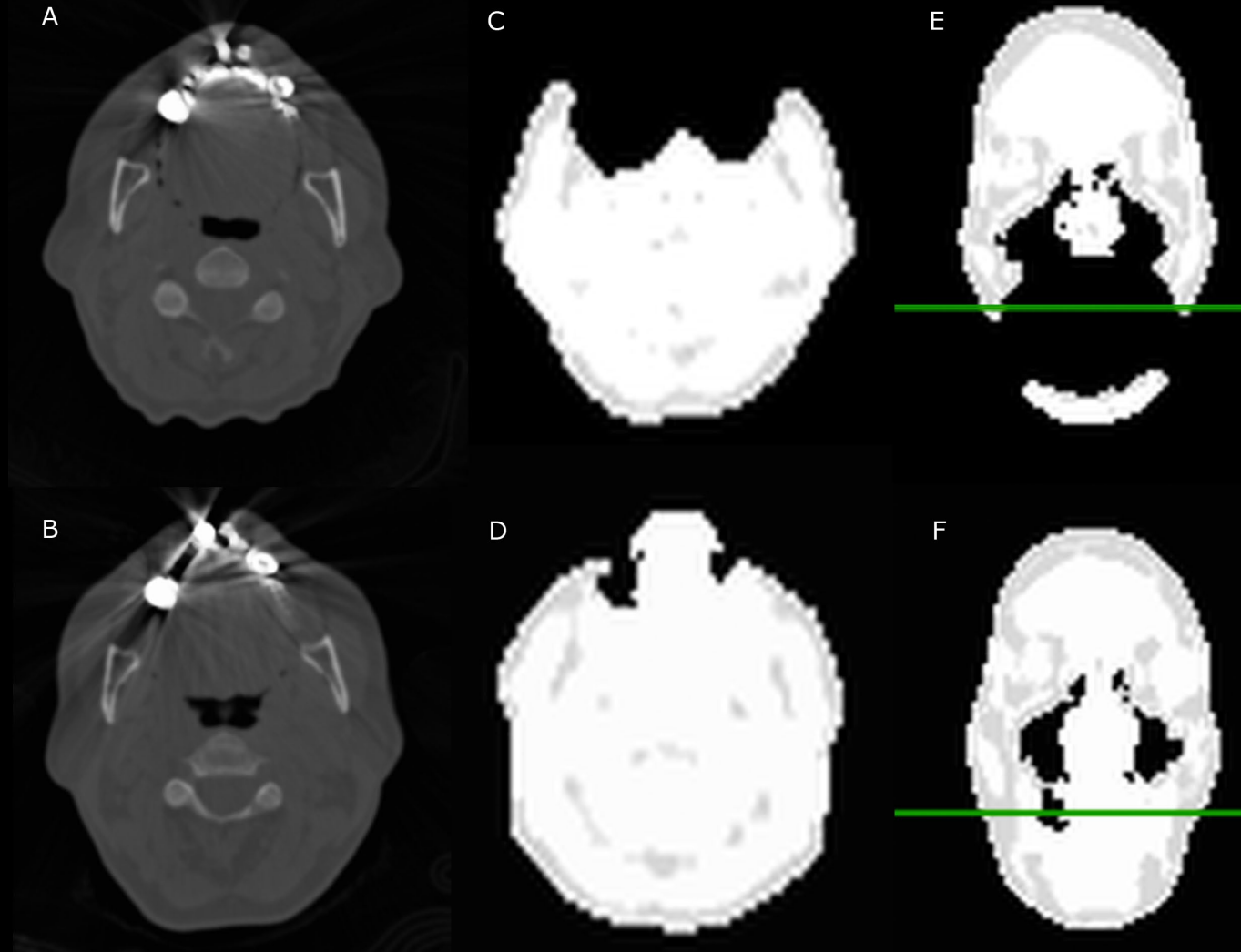


AC-PET_{MR}

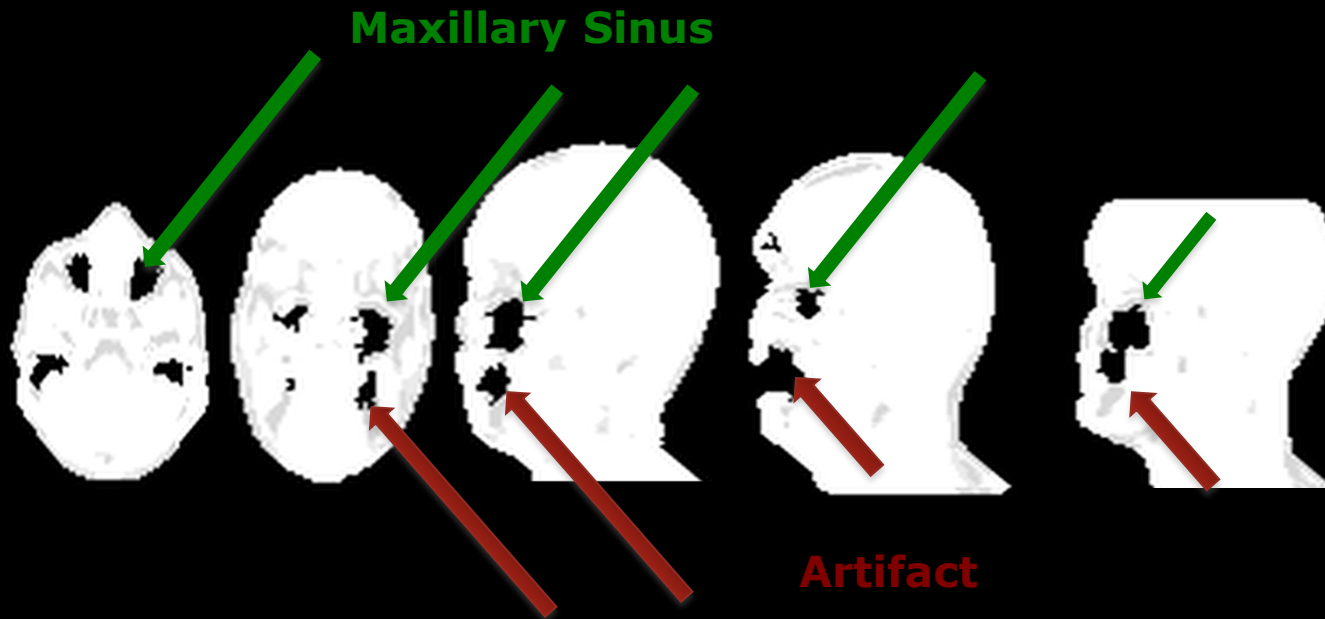


AC-PET_{MR}

Active Shape Models – and more



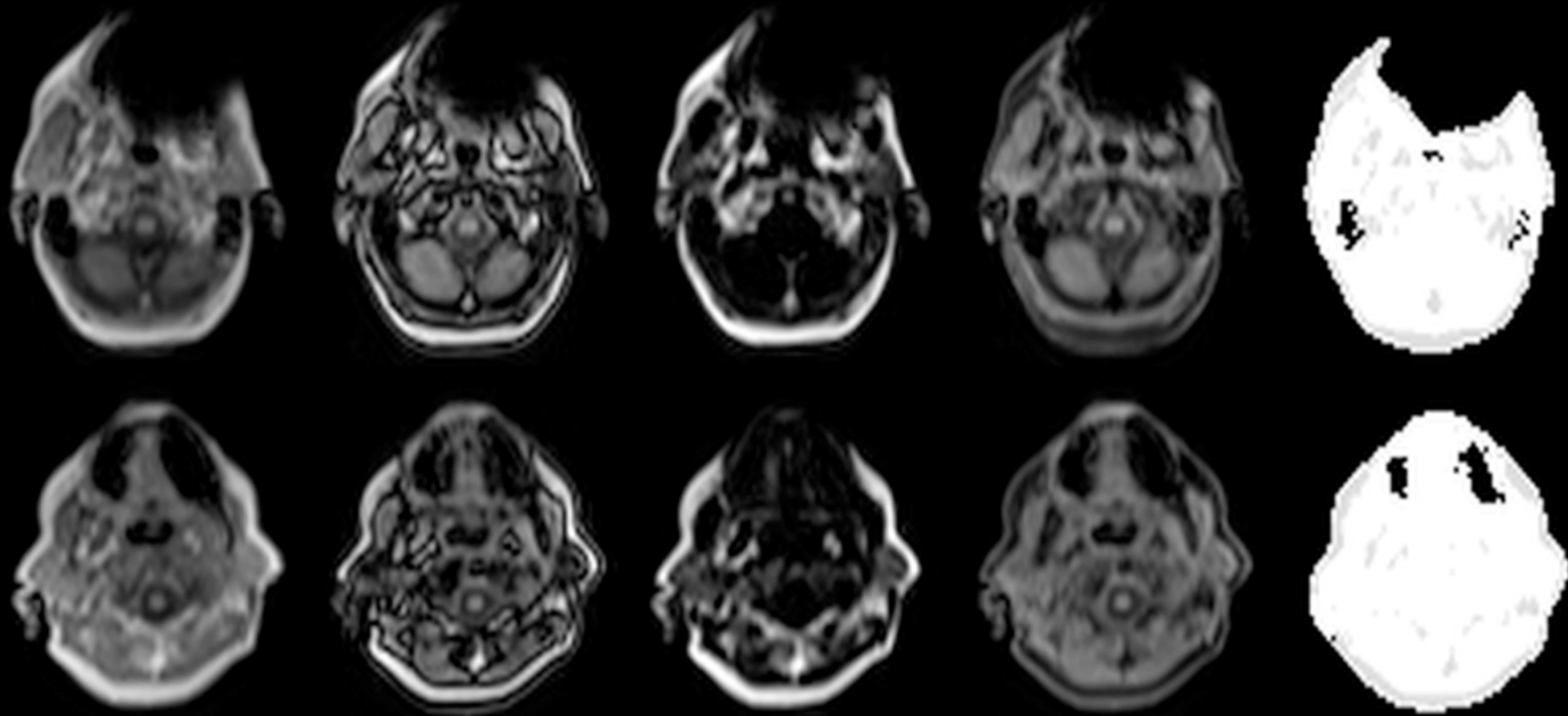
Active Shape Models – and more



Artifacts can be connected artificially with sinuses or background

Active Shape Models – and more

Outer holes = Signal voids breaching the anatomical surface



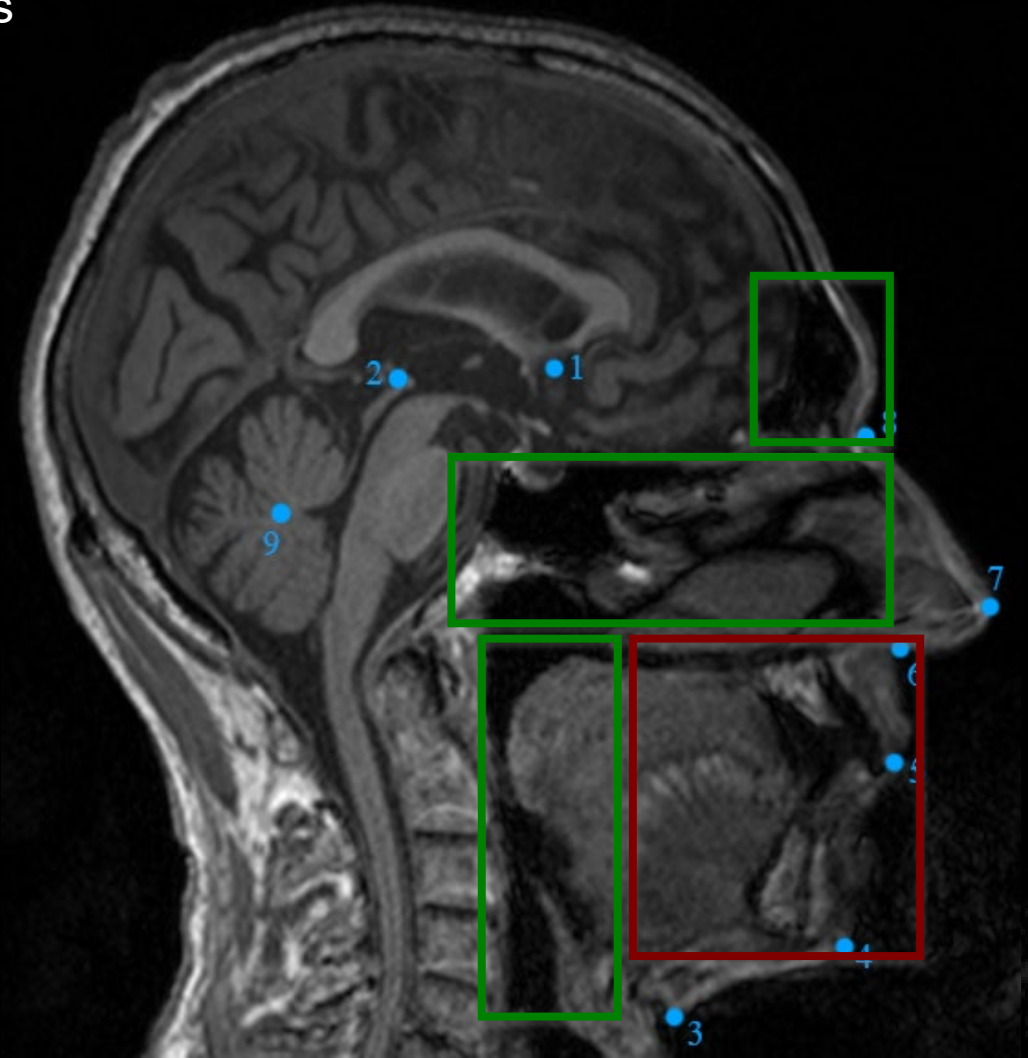
Inner holes = Signal voids within the anatomical surface

Active Shape Models – and more

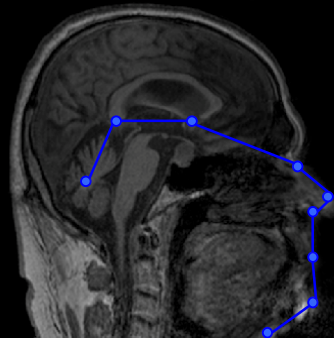
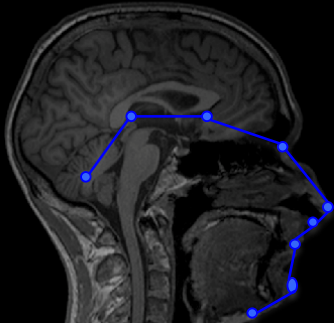
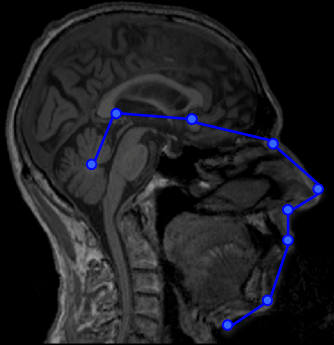
Artifacts can be separated from actual signal voids

How?

By the offset to a set of landmarks in 2D



Active Shape Models – and more



Procrustes analysis:

Transformation

$$X_i \rightarrow \rho_i X_i H_i + T_i$$

ρ : scaling

H : rotation

T : translation

Minimization problem

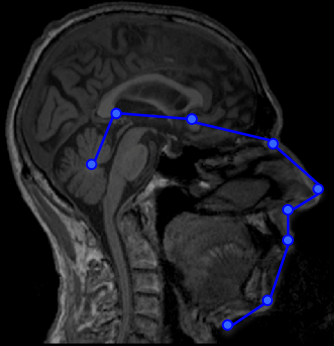
$$\sum_{i < s} \left\| (\rho_i X_i H_i + T_i) - (\rho_s X_s H_s + T_s) \right\|_F^2$$



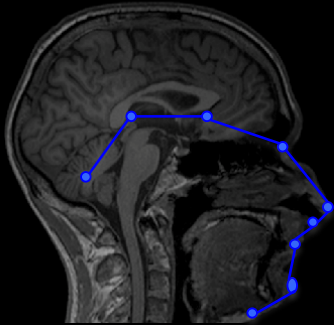
Landmarks on 7 patients

Active Shape Models – and more

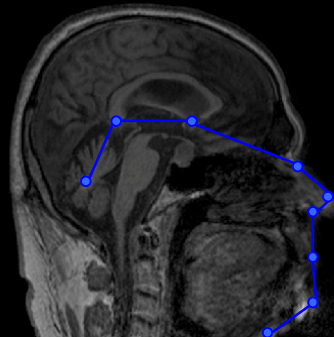
Eigenvalue	$\frac{\lambda_i}{\lambda_T} \times 100\%$
λ_1	41%
λ_2	25%
λ_3	19%
λ_4	8%
λ_5	5%
λ_6	2%



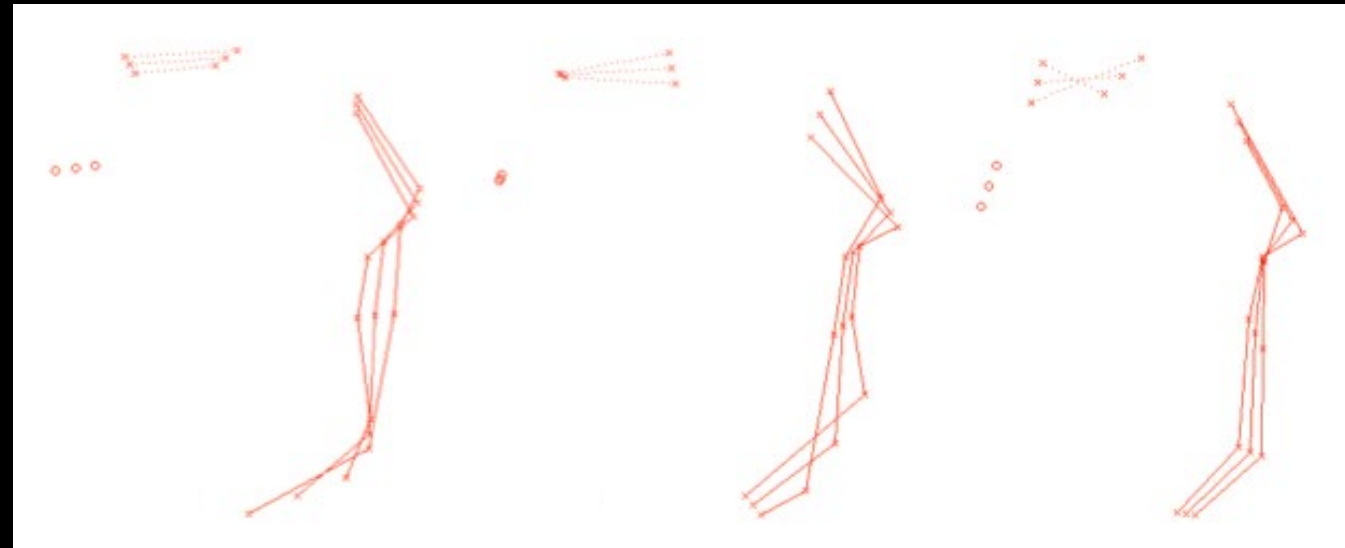
Mode 1: Mouth, horizontal & cerebellum



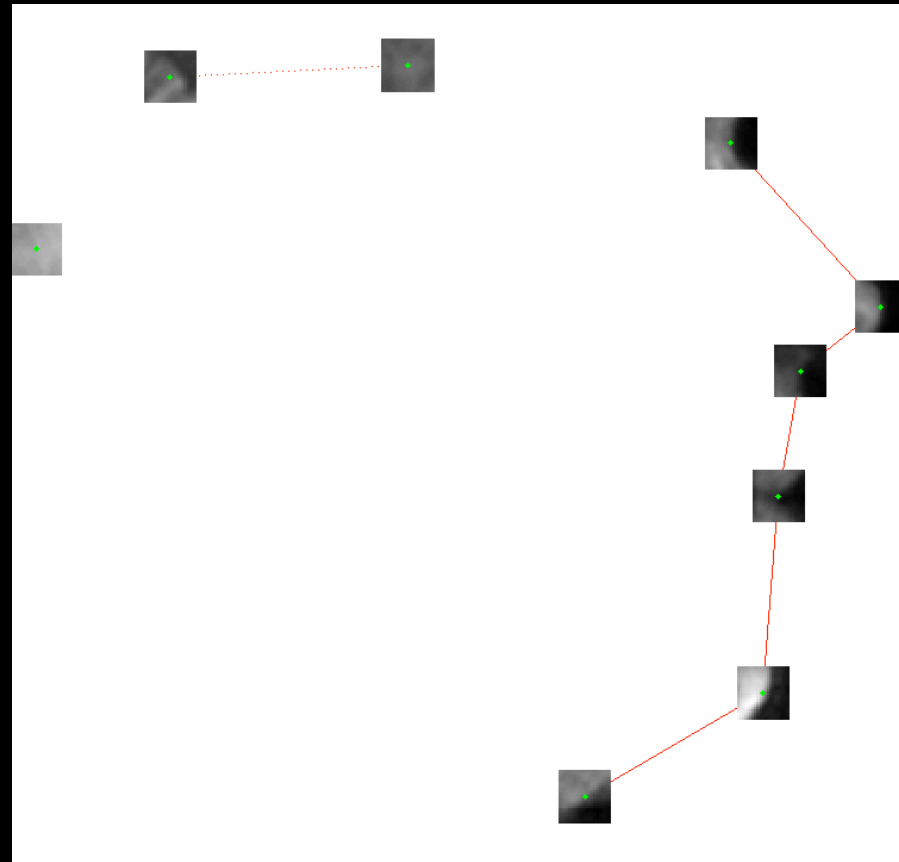
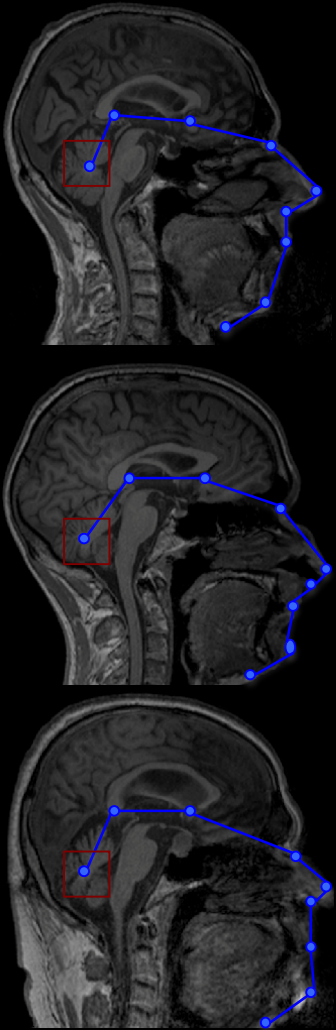
Mode 2: Chin



Mode 3: Anterior-posterior landmarks in respect to each other & cerebellum

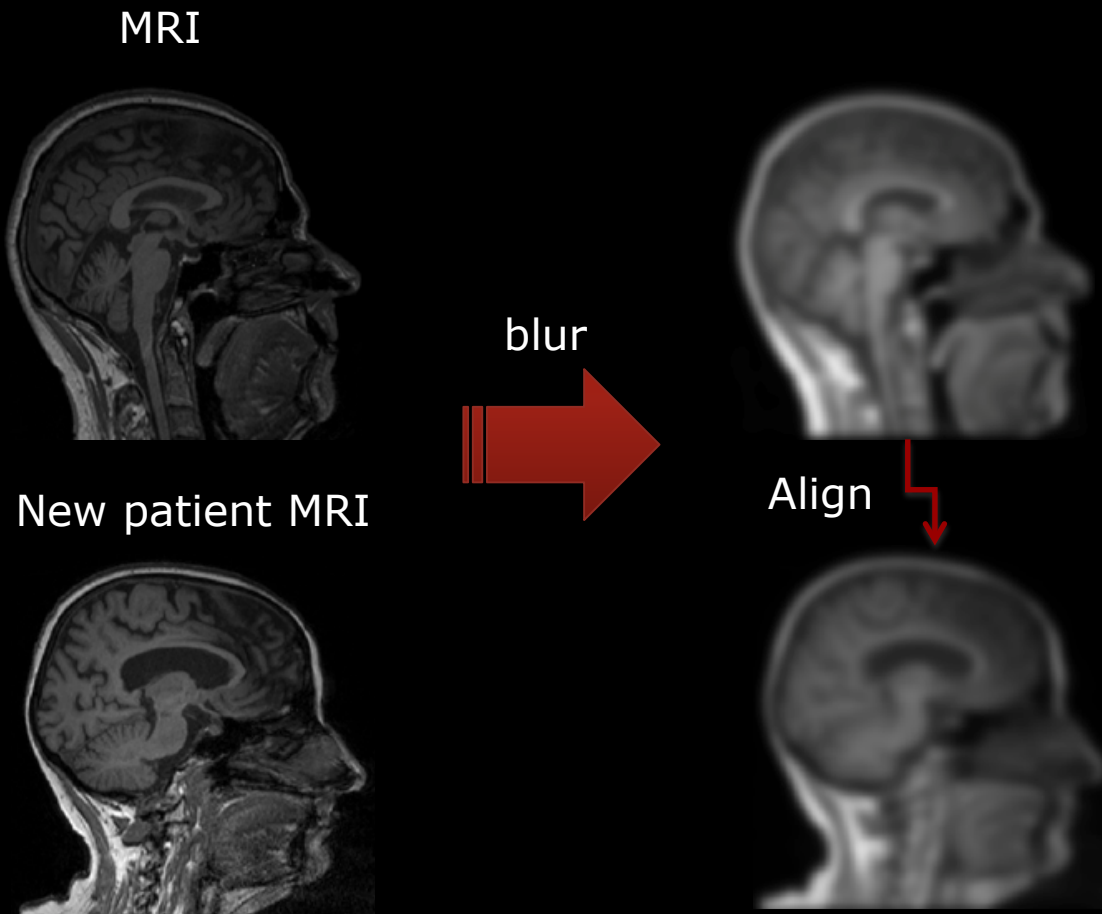


Active Shape Models – and more



Mean patches from 5 patients

Active Shape Models – and more



1. Blur MRI images from "atlas" patient and new patient
2. Align the two using rigid transformation
3. Apply the same transformation to the shape of the atlas patient

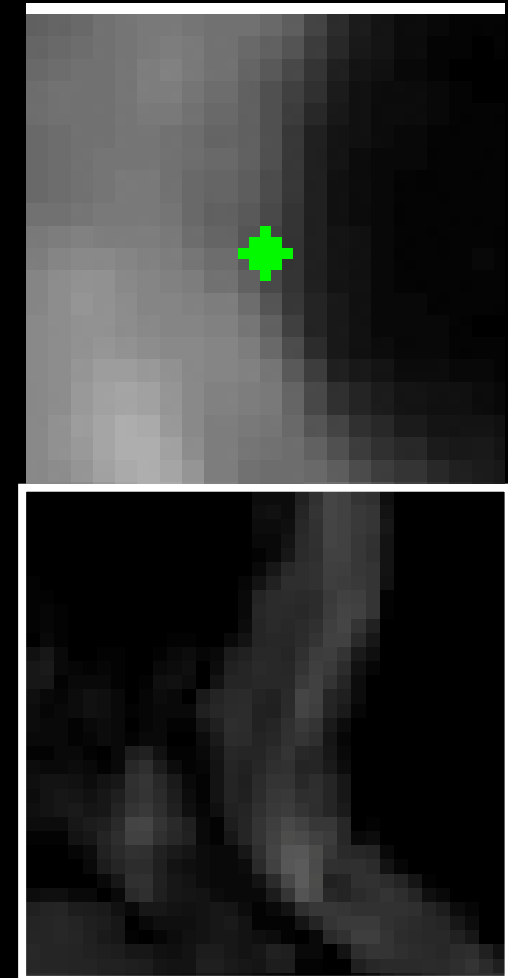
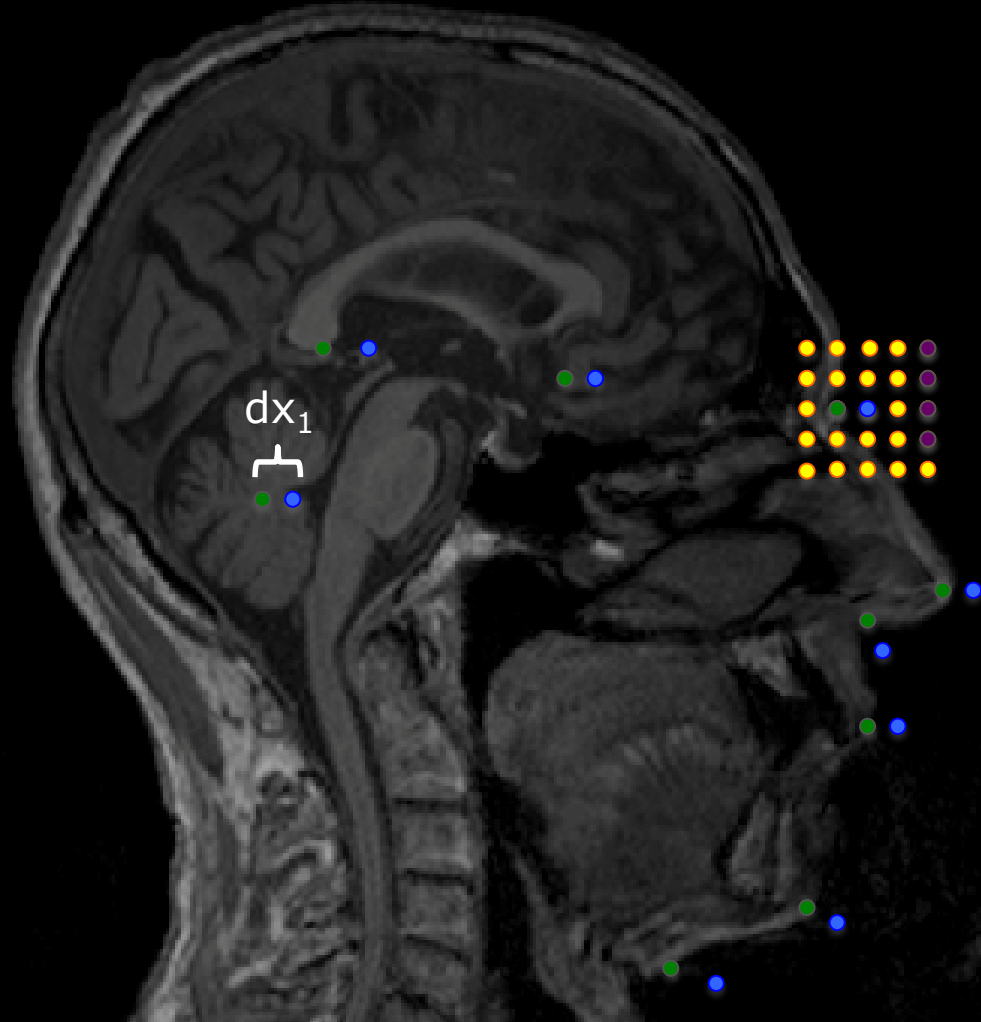
Active Shape Models – and more

Offset to
mean shape:

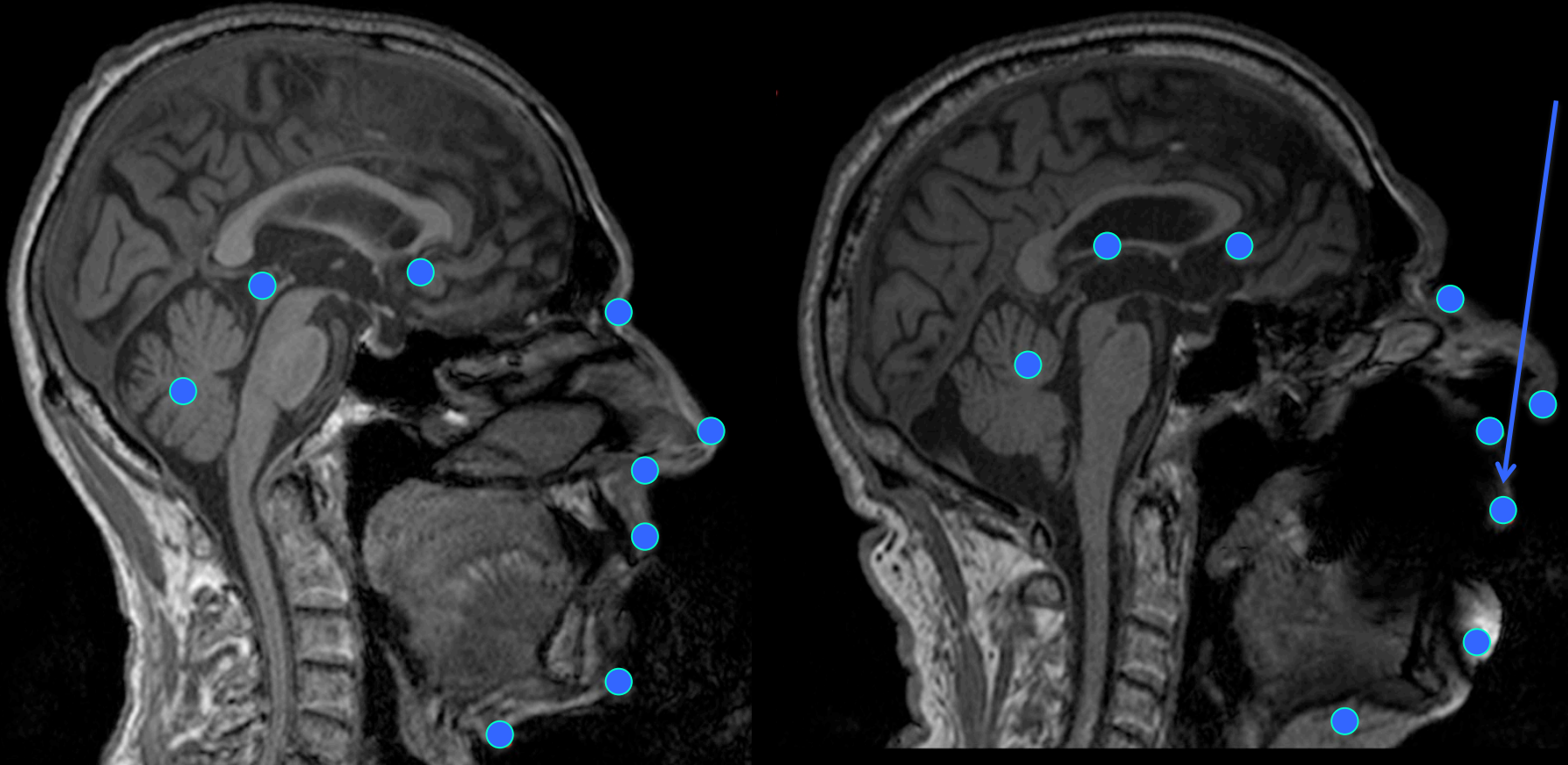
$$d\mathbf{x} = (dx_1, \dots, dx_n)$$

Projected to legal
shape space:

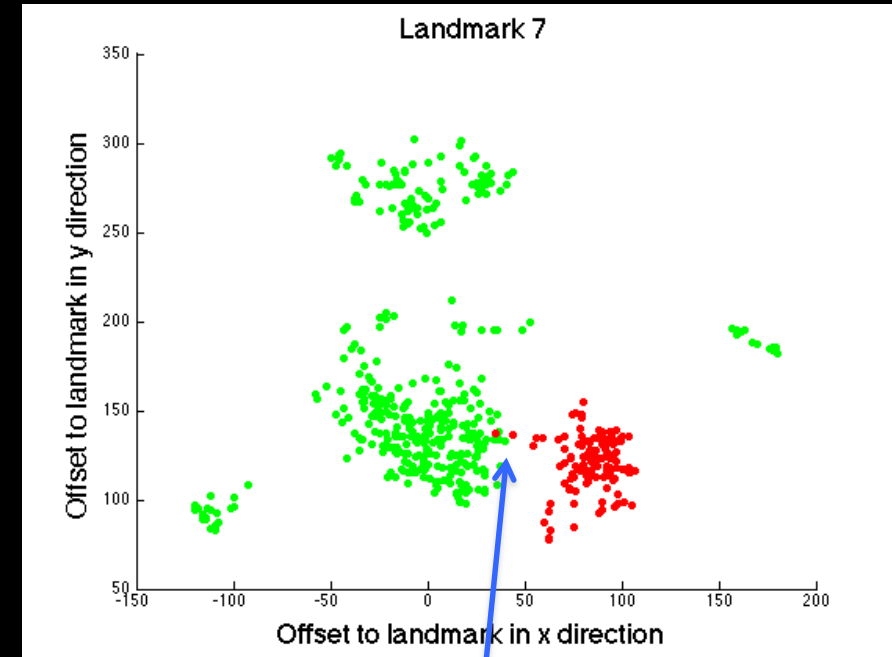
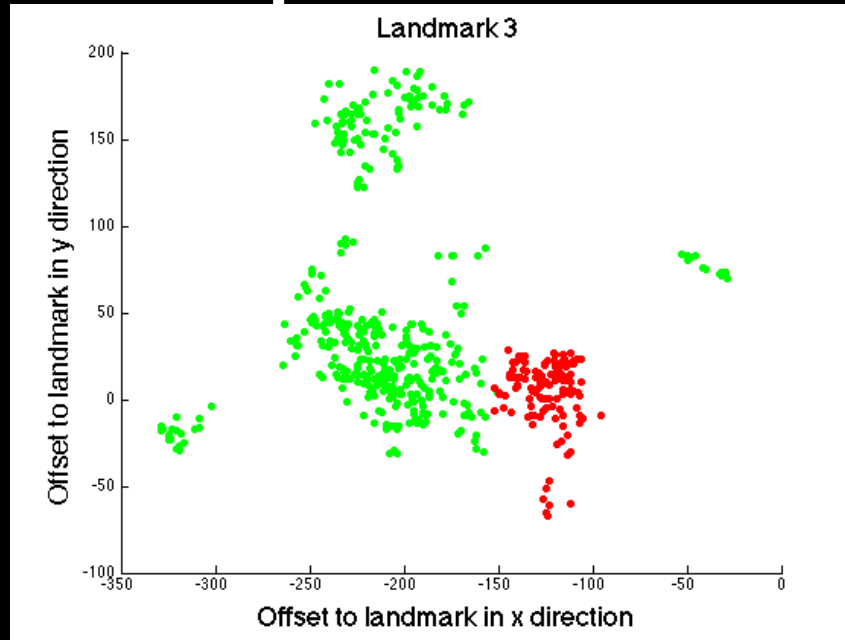
$$d\mathbf{y} = \phi^T d\mathbf{x}$$



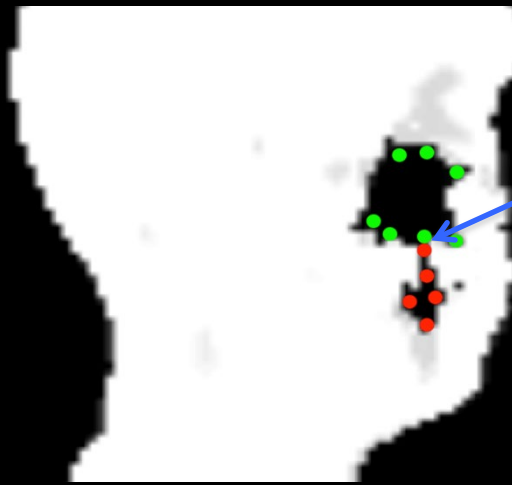
Active Shape Models – and more



Active Shape Models – and more



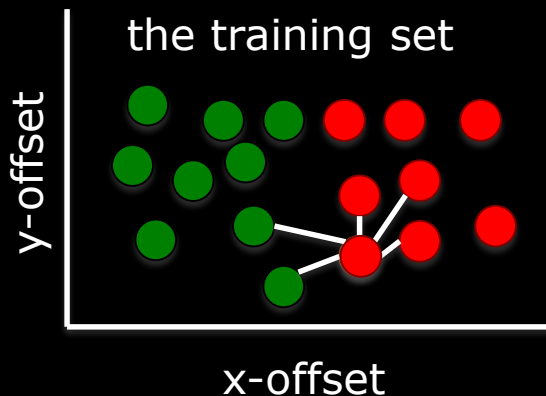
5 patients
650 non-artifact pixels
210 artifact pixels



Active Shape Models – and more



Offsets to a landmark in the training set

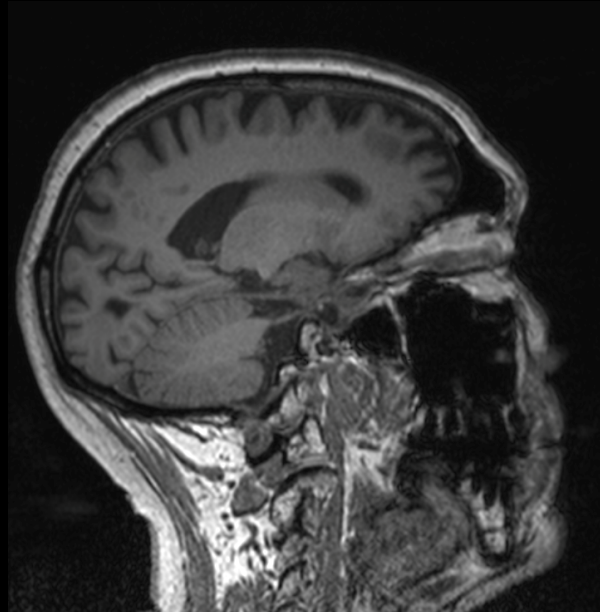


□ Classify using kNN

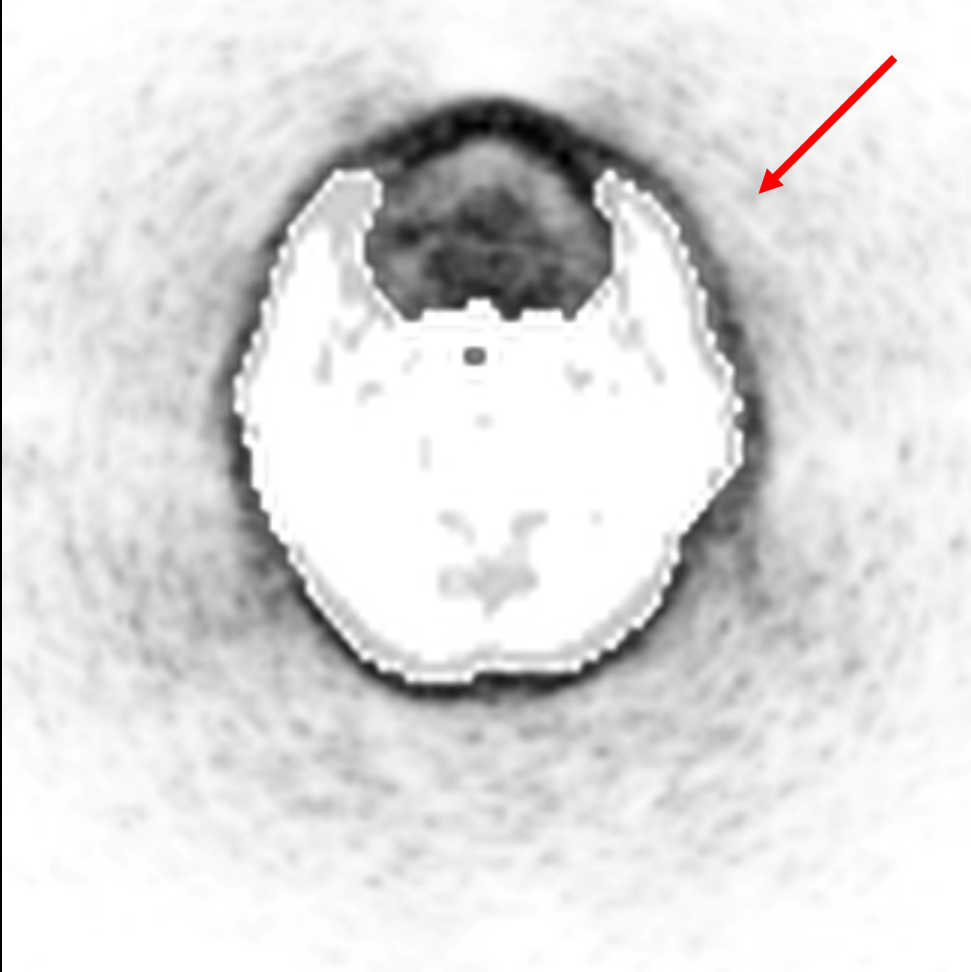
- For each pixel in a signal void
 - Find the offset to each landmark
 - Find 5-Nearest-Neighbors
 - Majority of neighbor-labels decides the landmark
 - Majority of landmark-labels decides the class

Classification: ●

Active Shape Models – and more



Active Contours: Chan Vese



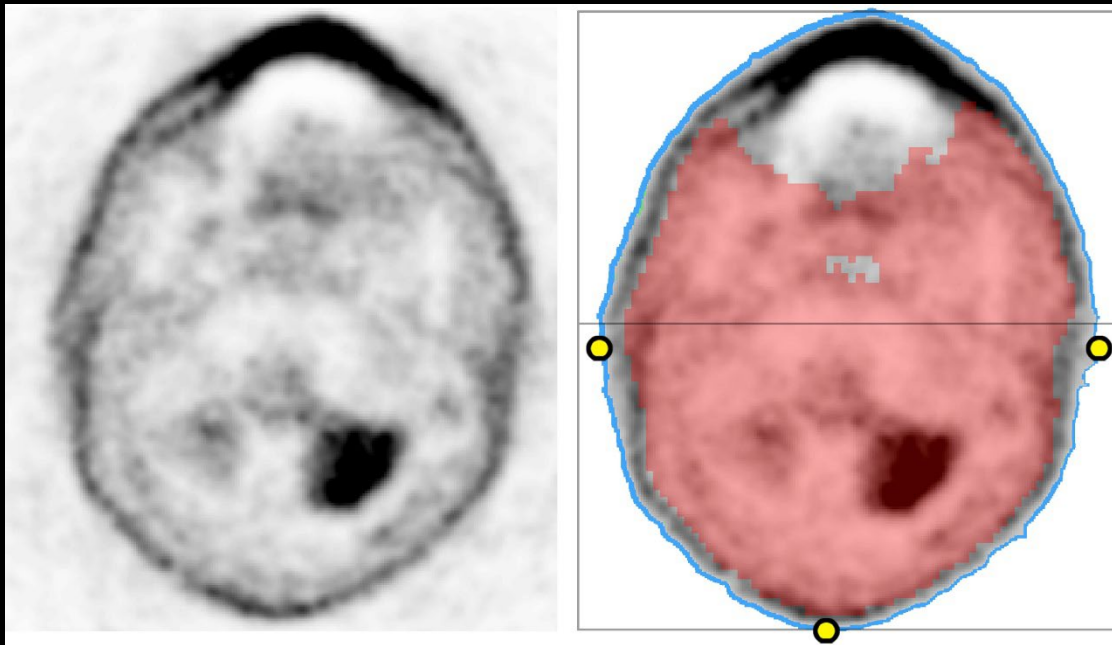
- "Outer holes" cannot be corrected easily by MRI
- NAC-PET holds information about outer contour
- ... but contains noise and needs to be delineated

Active Contours: Chan Vese

$$E_{CV}(\phi, c_i, c_o) = \mu \cdot \text{Length}(\phi) + \lambda_i \int_{\Omega} H(\phi) |u_0 - c_i|^2 + \lambda_o \int_{\Omega} H(-\phi) |u_0 - c_o|^2$$

Inside contour
Outside contour

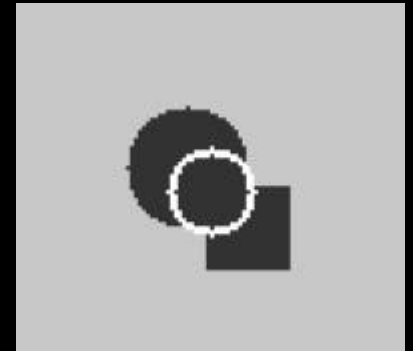
Length of contour
NAC-PET
Mean of areas



$F(\phi) > 0$
 $F(-\phi) \approx 0$



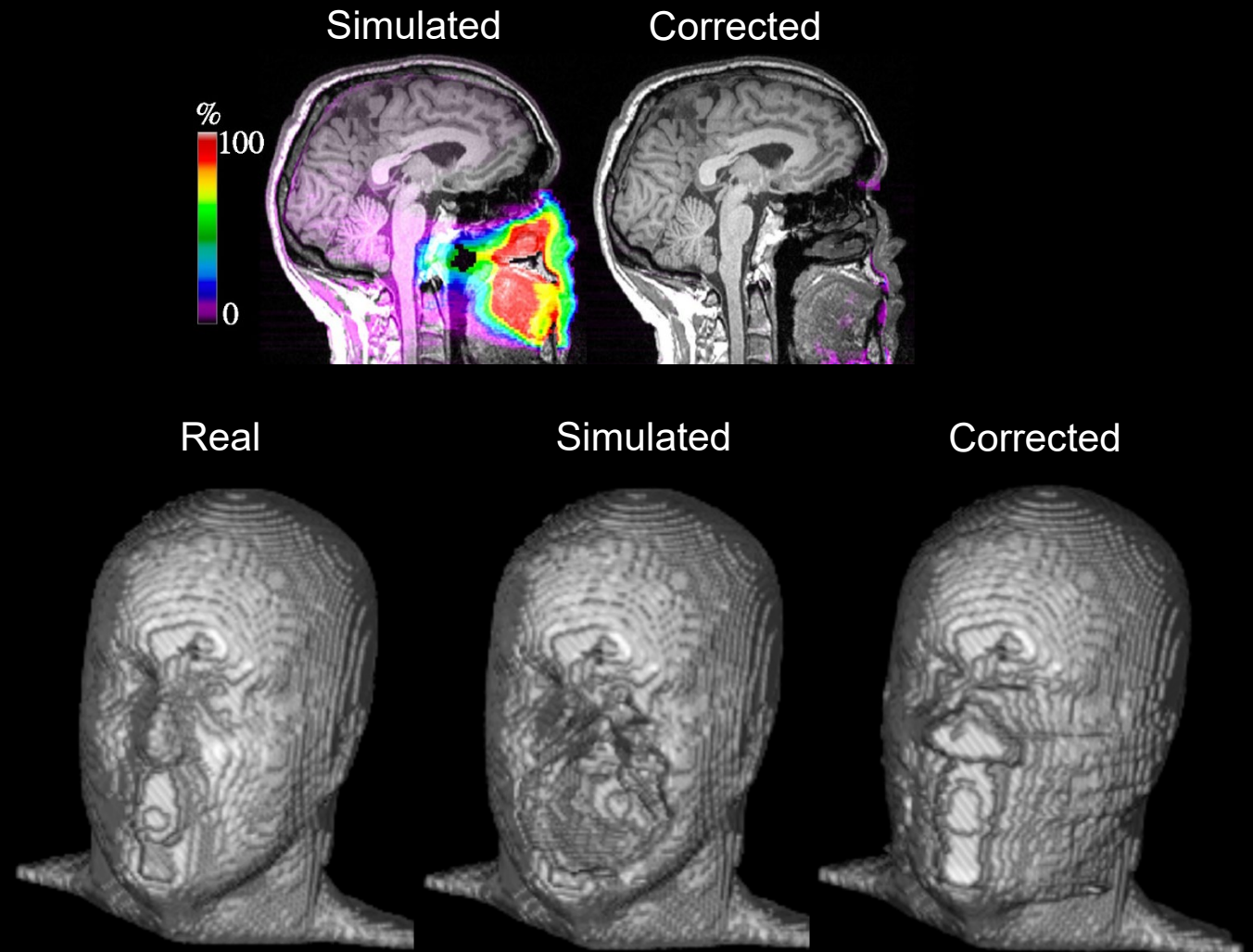
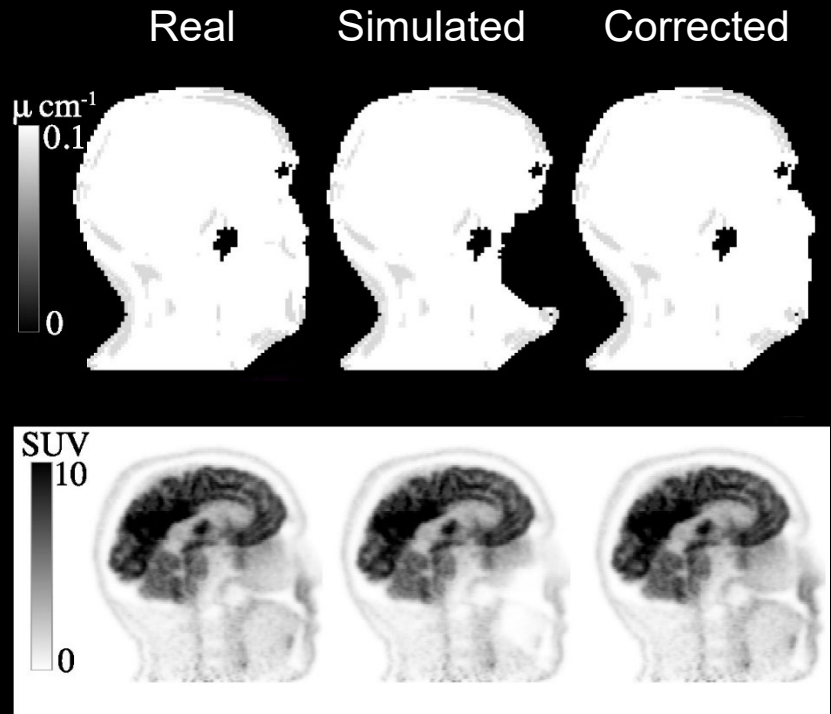
$F(\phi) \approx 0$
 $F(-\phi) > 0$



$F(\phi) \approx 0$
 $F(-\phi) \approx 0$



Active Contours: Chan Vese



What did you learn today

- Many of the topics taught during this course can be useful for image analysis at an imaging department in a hospital
- Topics like preprocessing are always used before any imaging project
- Registration are used to align scans within a patient examination, and across examinations
- Simple tools are often wanted as it
 - Works well with limited data
 - Strengthens the explainability of a method

