

# Image Enhancement and Preprocessing continued

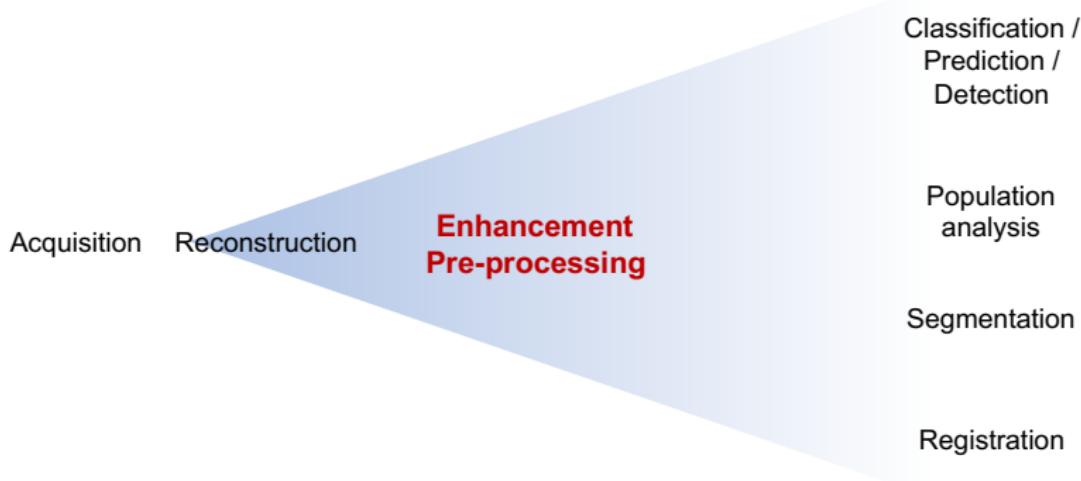
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ETH Zürich

March 03, 2020

## Section 1

### Introduction



Enhancement and pre-processing tasks form initial steps for any type of analysis. Same applies to methods that use machine learning, probabilistic, variational or energy-based formulations.

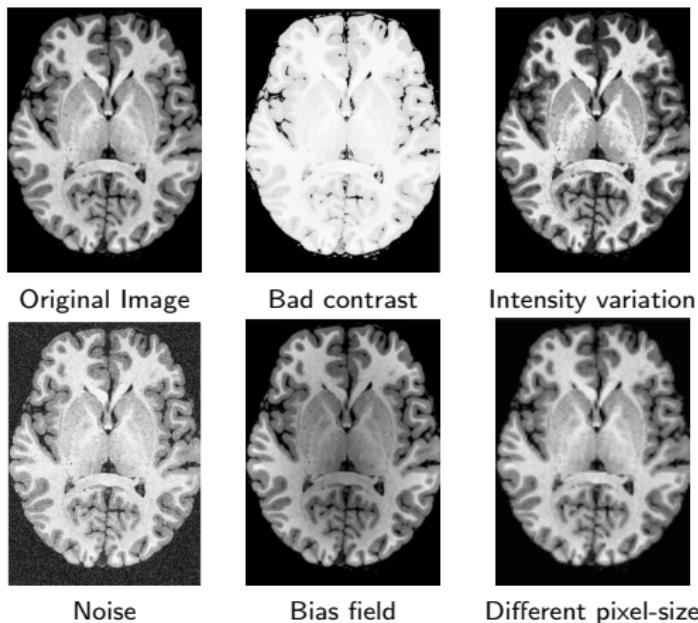
# Image enhancement and pre-processing

Enhancement of four conditions

- Bad contrast
- Varying intensity statistics
- Noise
- Bias field
- Variation in pixel-size

Why enhancement?

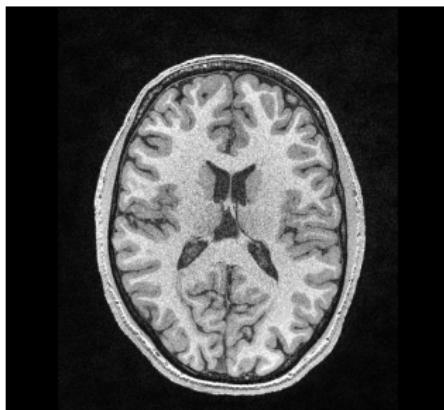
- simplifying interpretation
- better visualization
- normalization for further processing



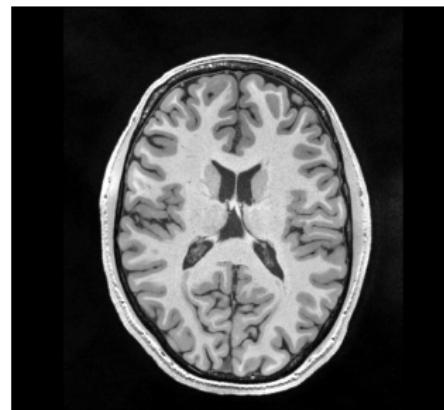
## Outline

- Contrast enhancement
- Intensity normalization
- Noise suppression
- Exercise / Challenge
- Bias correction
- Variations in image and pixel-size

## Noise suppression challenge: Discussion



Single acquisition



Average of 8

### Challenge

High-resolution MRI is hard to acquire. One way is to acquire many images of the same subject and average them. 8 or so images yield good SNR but this leads to long acquisitions. Can you reduce this time with noise suppression?

More details in the moodle platform.

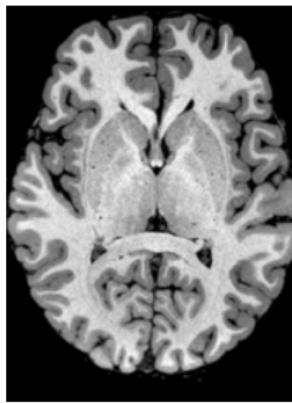
## Section 2

### Bias Correction

## Outline

- Contrast enhancement
- Intensity normalization
- Noise suppression
- Exercise / Challenge
- Bias correction
  - What is bias field?
  - N3 algorithm
- Variations in image and pixel-size

## What is bias field?



Normal image

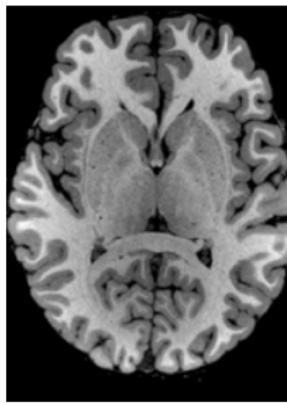
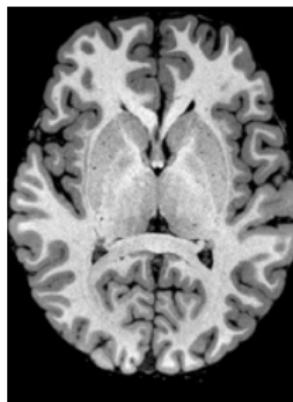


Image with bias

### ■ Acquisition artifact

- Field inhomogeneity
- Electrical characteristics of the tissue
- Poor uniformity of coils
- Gradients and eddy currents

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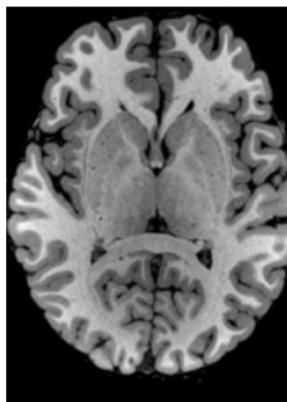
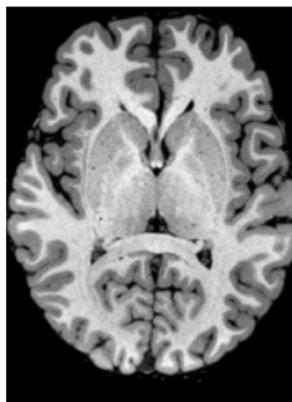


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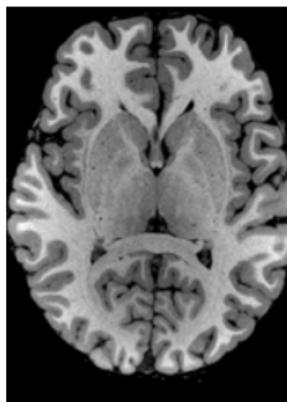
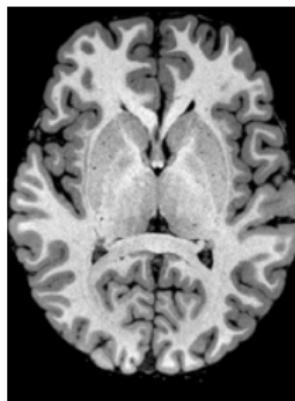


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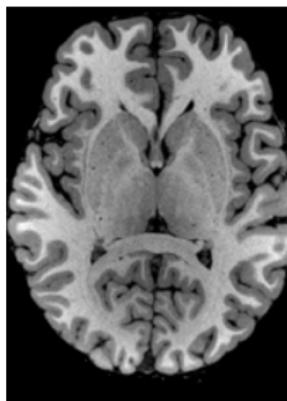
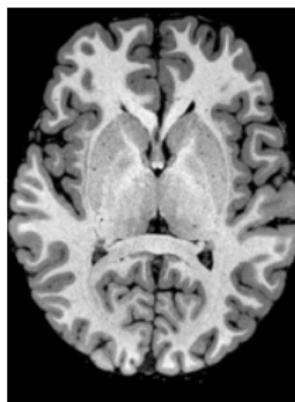


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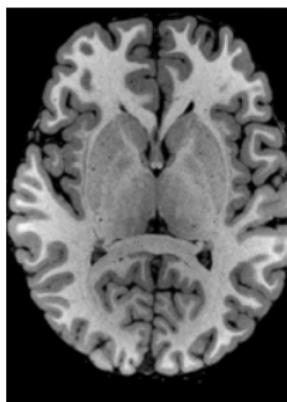
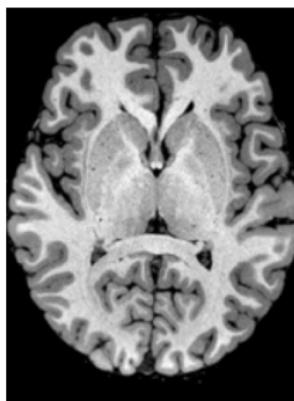


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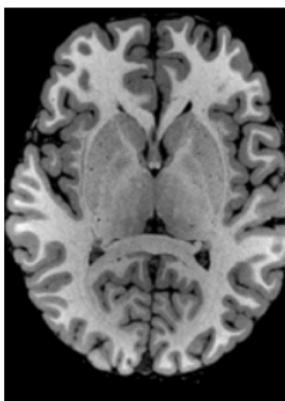
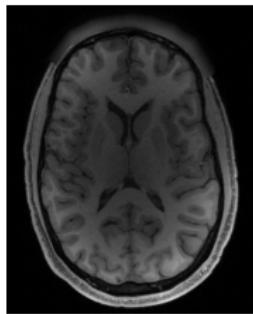


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- Non-linear effect on intensities that vary spatially
- Remove it as a pre-processing step

What does it look like?



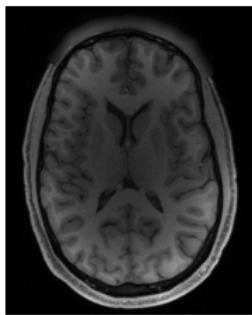
Image



Bias field

## What does it look like?

- Smoothly varying

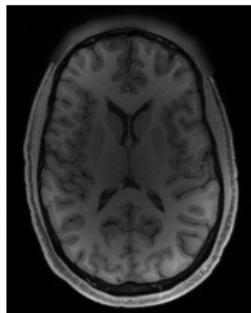


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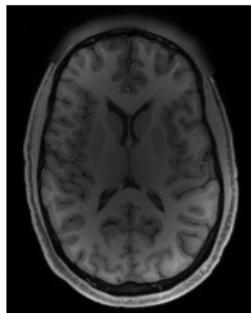
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Bias field

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- Does not depend on the underlying tissue to the first order approximation

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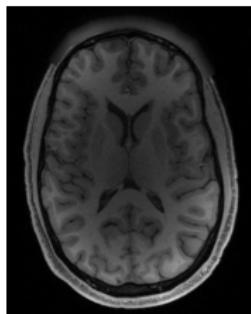
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## What does it look like?



Image



Bias field

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- In reality it also depends on the tissue but this effect is mostly ignored
- Approximate mathematical model:

$$j = bi + n,$$

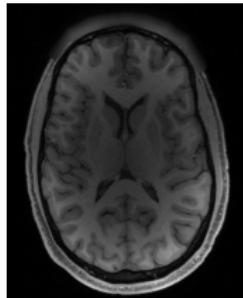
$i$ : real image,  
 $b$ : bias field,  
 $n$ : noise and  
 $j$ : observed image

- Multiplicative factor

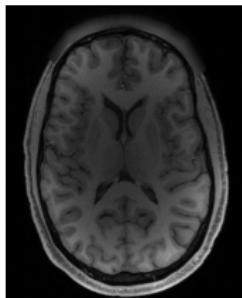
└ Bias Correction

└ What is bias field?

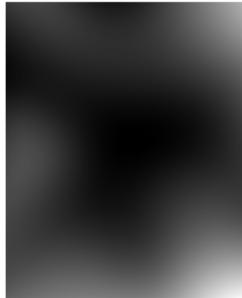
## Basic methods to correct it



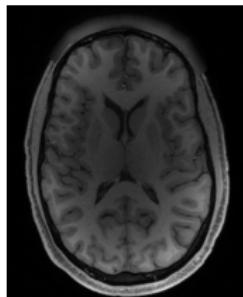
## Basic methods to correct it



- At the hardware side with extra measurements
  - Ideally a perfect solution
  - Additional measurements makes it hard for retrospective data analysis and usually not integrated
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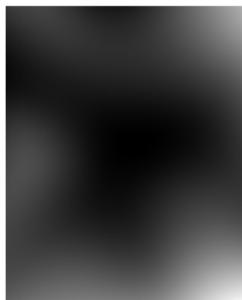
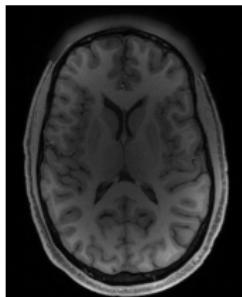
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  - Voxels expected to have similar intensity
  - Human in the loop - a very good solution
  - Need for human interaction

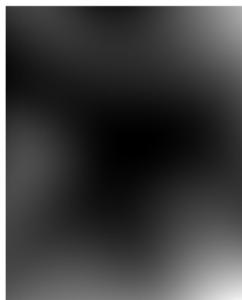
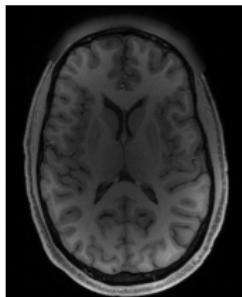


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- N3 algorithm
  - Generic
  - Most popular solution
  - [Sled, Zijdebos and Evans, IEEE TMI 1998]
  - We will study N3 in this lecture.

## N3 algorithm - principle

Starting from the main model

$$\tilde{j}(x) = \tilde{b}(x)\tilde{i}(x) + \tilde{n}(x)$$

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### Key point

N3 analyzes the distribution of  $j$ ,  $b$  and  $i$  to understand the effect of the bias term to understand the effects of the bias term.

### Assumption

N3 assumes independence between pixels when considering distributions of  $j$ ,  $b$  and  $i$ .

$j(x) = b(x) + i(x)$  is a sum of random variables. Let's study how distribution of  $j$  is related to the distribution of  $i$  and  $b$ .

## Sums of random variables

Let  $a$  and  $b$  be two independent random variables with pdfs denoted by  $f_a(a)$  and  $f_b(b)$ . Let us define another random variable

$$c = a + b$$

Can we compute the distribution of  $c$ , i.e.  $f_c(c)$ , using distributions of  $a$  and  $b$ ?

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The CDF is given as

$$F_c(c) = \int_{-\infty}^{\infty} \int_{\infty}^{c-b'} f_a(a') f_b(b') da' db'.$$

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### Convolution

$c = a + b \Rightarrow f_c = f_a * f_b$  when  $a$  and  $b$  are independent random variables.

## Back to bias problem

$$j(x) = b(x) + i(x),$$

Denote the probability distributions of  $j$ ,  $b$  and  $i$  with  $J(j)$ ,  $B(b)$  and  $I(i)$ . The probability distribution  $J$  is the following convolution:

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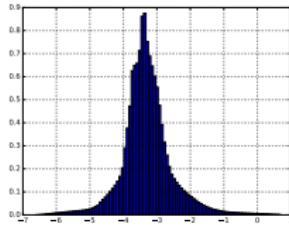
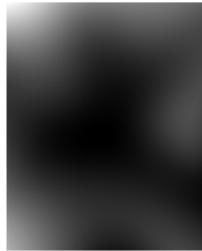
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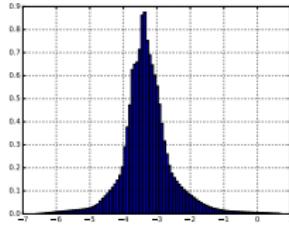
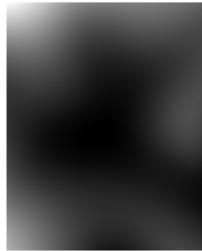
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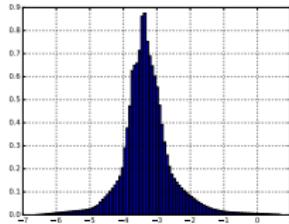
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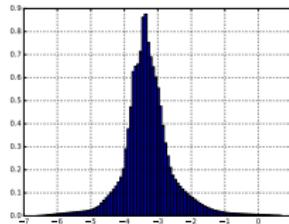
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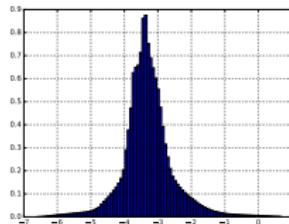
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- $J$  is a smoothed version of  $I$

## Two components in the problem

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The estimation must determine  $I(i)$  and  $b$  given only  $j$ . We divide this into two parts:

- Field estimation: Estimation of  $b$  given  $I(i)$ .
- $I(i)$  estimation: Estimation of  $I(i)$ .

Always assuming  $B$  is a Gaussian with small standard deviation.

## Component 1: Field estimation

One component is field estimation. If we are given the distribution  $I(i)$  we can estimate the bias field at any point by

$$\mathbb{E}[i|j] = \int_{-\infty}^{\infty} ip(i|j)di$$

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$$\begin{aligned}\mathbb{E}[i|j] &= \int_{-\infty}^{\infty} i \frac{p(i,j)}{p(j)} di \\ &= \frac{1}{J(j)} \int_{-\infty}^{\infty} ip(i,j)di, \quad b = j - i \\ &= \frac{1}{J(j)} \int_{-\infty}^{\infty} ip(i)p(b)di \\ &= \frac{\int_{-\infty}^{\infty} iB(j-i)I(i)di}{\int_{-\infty}^{\infty} B(j-i)I(i)di}\end{aligned}$$

## Component 1: Field estimation

One component is field estimation. If we are given the distribution  $I(i)$  we can estimate the bias field at any point by

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We know all the terms in this equation and  $i$  is one dimensional, so, we can solve it numerically with ease.

## Component 2: $I(i)$ estimation

### Key observation

$J(j)$  is a smoothed version of  $I(i)$ , and  $B(b)$  is roughly Gaussian with a small standard deviation.

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$$\begin{aligned} \text{Deblurring filter: } \hat{F} &= \frac{\mathcal{F}\{B\}^*}{|\mathcal{F}\{B\}|^2 + Z^2} \\ \mathcal{F}\{I\} &\approx \hat{F}\mathcal{F}\{J\} \end{aligned}$$

with  $Z$  being a small number and  $\mathcal{F}\{B\}^*$  denotes the complex conjugate.

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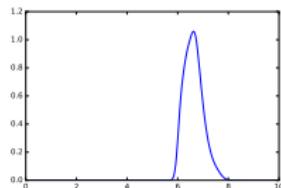
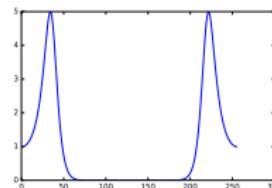
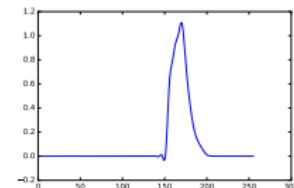
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**Note:** This is inverse filtering!

## Intermediate summary

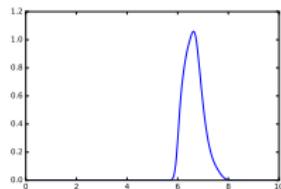
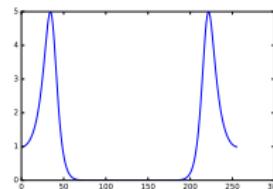
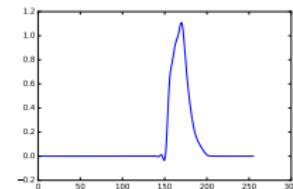
- Estimate of the unbiased image's distribution:  $I(i) \approx \mathcal{F}^{-1}(\hat{\mathcal{F}}\mathcal{F}\{J\})$

 $J(j)$  $|\hat{F}|$  $I \approx \mathcal{F}^{-1}(\hat{\mathcal{F}}\mathcal{F}\{J\})$ 

- field estimate:  $b_e(j) = j - \frac{\int_{-\infty}^{\infty} iB(j-i)I(i)di}{\int_{-\infty}^{\infty} B(j-i)I(i)di}$
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- But there are two things...

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- Solution: Smooth approximation based on anchor points using Splines or Radial Basis Function approximation:

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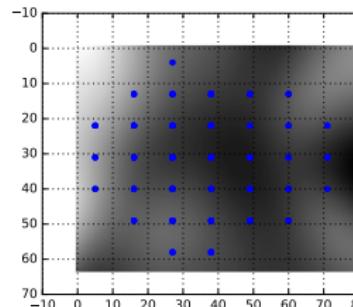
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- Example



## 2. Need for iterations

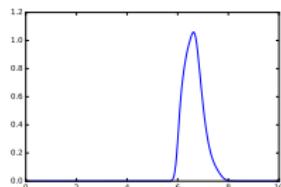
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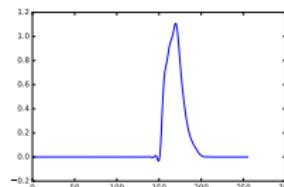
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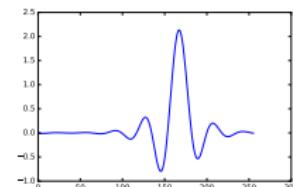
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$J(j)$



$\text{std} = 0.1$

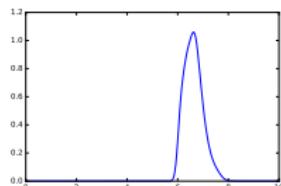
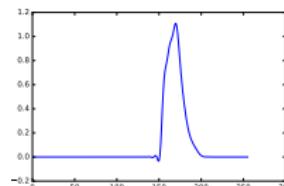
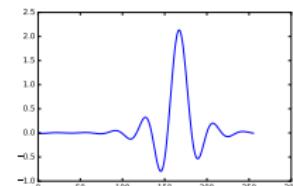


with  $\text{std} = 0.5$

Remember they have to be distributions ( $> 0$ ). **Deconvolution is hard.**

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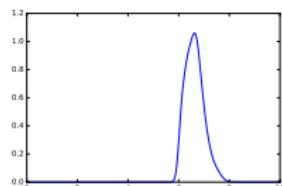
 $J(j)$  $\text{std} = 0.1$ with  $\text{std} = 0.5$ 

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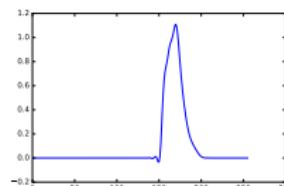
- Solution: Iterate many times with small Gaussians (small stds)

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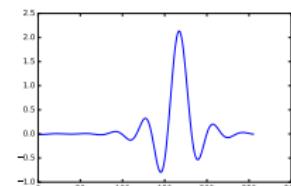
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$J(j)$



$\text{std} = 0.1$



$\text{with std} = 0.5$

Remember they have to be distributions ( $> 0$ ). **Deconvolution is hard.**

- Solution: Iterate many times with small Gaussians (small stds)
- Not a bad approximation: large Gaussians are convolutions of smaller ones

## Algorithm

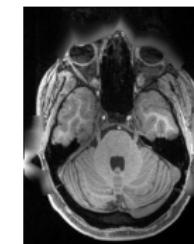
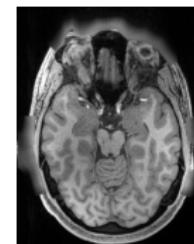
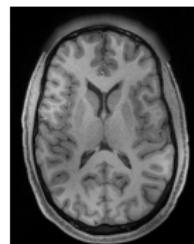
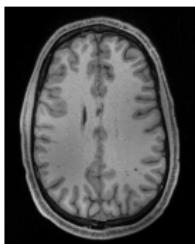
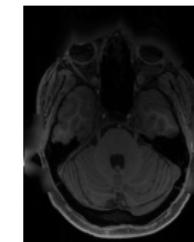
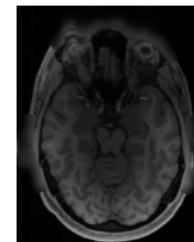
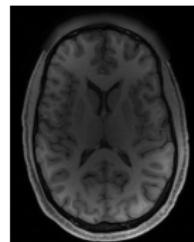
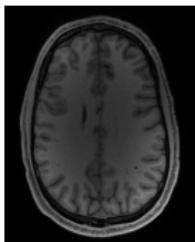
---

**Algorithm 1** Bias correction with N3

---

- 1:  $b = 0, j = \log(\tilde{j})$  observed image
  - 2:  $\hat{F} = \frac{\hat{B}^*}{|\hat{B}|^2 + Z^2}$
  - 3: **while**  $\|\Delta b\| < \text{eps}$  **do**
  - 4:   Estimate  $J(j)$  with kernel density estimation or simple histogram
  - 5:   Compute  $I(i) = \mathcal{F}^{-1}(\hat{F}\mathcal{F}\{J\})$
  - 6:   Compute  $\Delta b_e = j - \mathbb{E}(i|j)$  at  $x_n$  anchor points.
  - 7:   Get the smooth estimate  $\Delta b(x) = \sum_{n=1}^N \Delta b_e(x_n)K(\|x - x_n\|)$  for all voxels
  - 8:   Update  $b = b + \Delta b$
  - 9:   Update  $j = j - \Delta b$
  - 10: **end while**
-

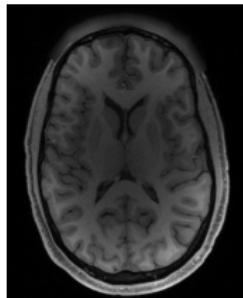
## Examples



## Analysis

- Bias correction with N3 is routinely used for MRI today
- There are variations available, e.g. N4ITK
- In the method there are multiple approximations
- Multiresolution solutions exist and perform better
- May need to run the tool a few times to get better results

## Question - Group discussion



- Manually placed landmarks
  - Voxels expected to have similar intensity
  - Human in the loop - a very good solution
  - Need for human interaction
- How would this exactly work?
- Let's only concentrate on structural brain MRI, e.g. like the images on the left hand-side.

## Section 3

Variations in image and pixel-size

## Outline

- Contrast enhancement
- Intensity normalization
- Noise suppression
- Exercise / Challenge
- Bias correction
- Variations in image and pixel-size
  - What is the problem?
  - How to tackle it.
  - How **not** to tackle it.

## Different image and pixel size and FOV



Fine pixel-size, small FOV



Coarse pixel-size, large FOV

### What is it?

- Images may come with different pixel and/or image size.
- They may have different Field-of-View (FOV).
- You may need to compare such “heterogeneous” images.

### Why does it happen?

- Differences in acquisitions
- in FOV
- in pixel size

## In which applications does it matter?

- Comparing measurements
- Registration
- Machine learning applications

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- Comparing measurements
  - measurements taken in an image have a correspondence in real life.
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  - comparisons need to take this into account.
  - e.g. longitudinal analysis, population statistics, normative distributions
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  - aligning images requires building point correspondence
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- parameters of a model are learned from a set of images.
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- Machine learning applications

- parameters of a model are learned from a set of images.
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- Effectively important for all applications

## How do we tackle this?

### Images of the same person one year apart

Sizes of the images on the slides are proportional to their number of pixels.

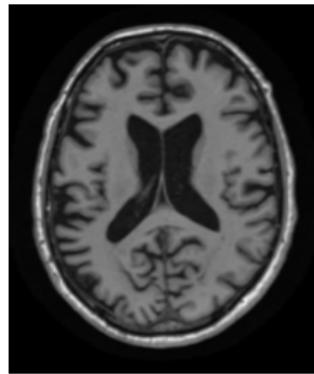


Image size: [454, 384]  
Pixel size: [1.25, 1.25]

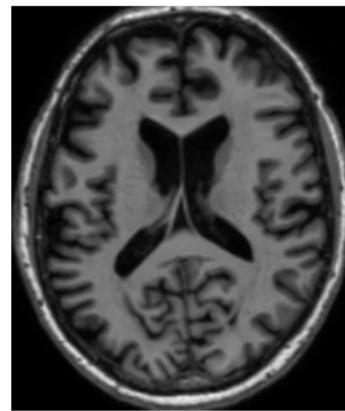


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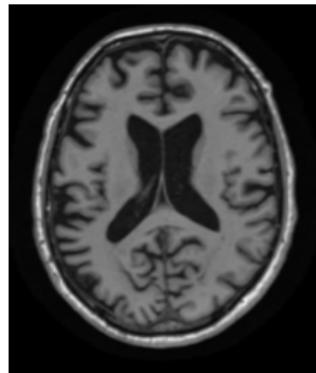


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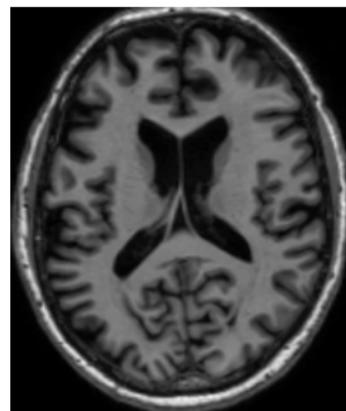


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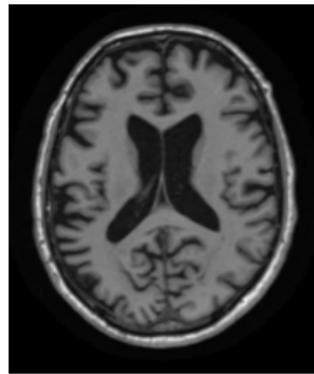


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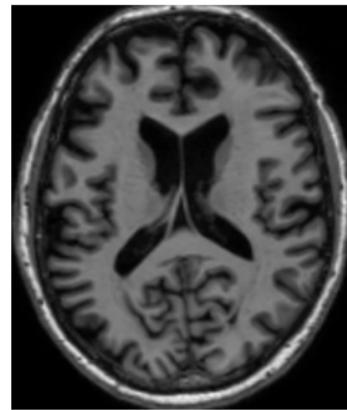


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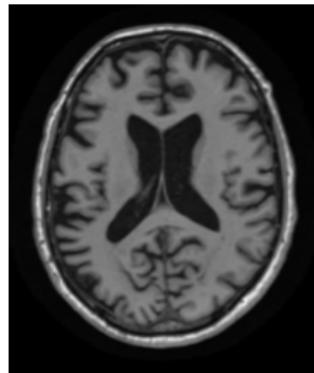


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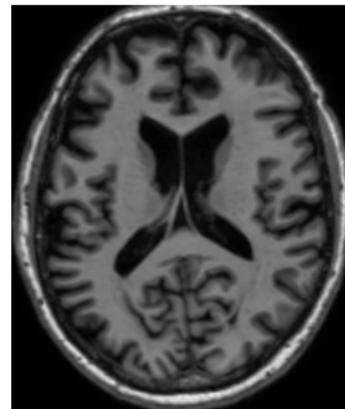


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## Match the pixel size

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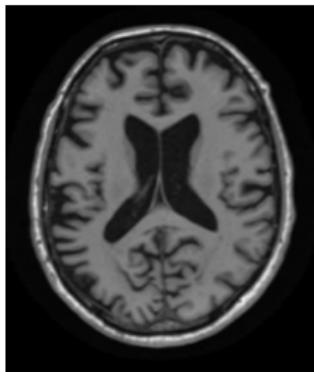


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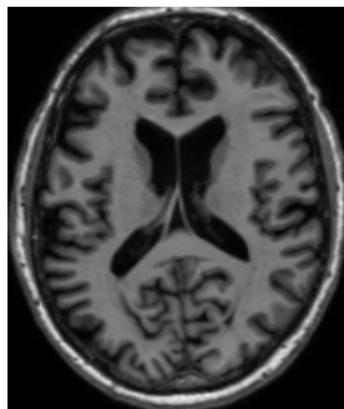


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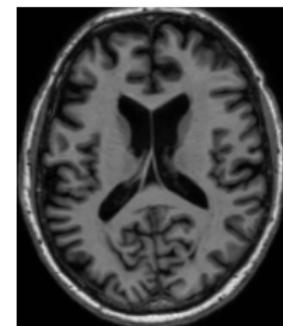


Image size: [395, 339]  
Pixel size: [1.25, 1.25]

- Resample one image to match the *pixel-size* of the other.
- Resize an image with a factor determined by the pixel-size. One example tool in python “scipy.ndimage.zoom”.

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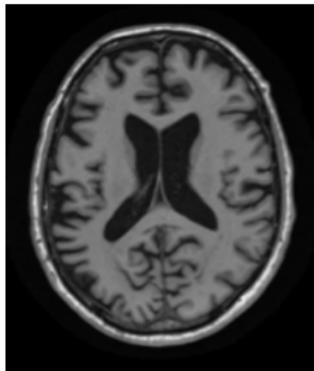


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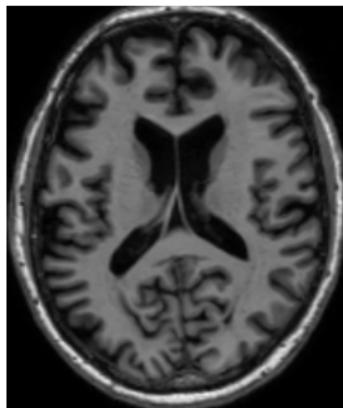


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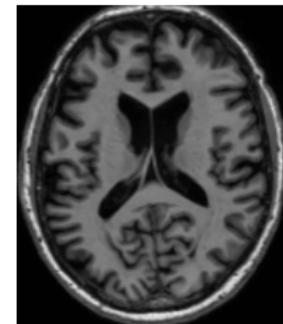


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- Resize an image with a factor determined by the pixel-size. One example tool in python “scipy.ndimage.zoom”.
- If volumetric images in 3D, then match the *voxel-size*.

## Match the pixel size

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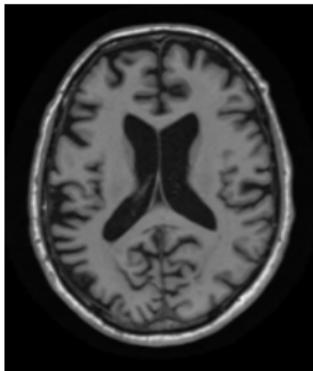


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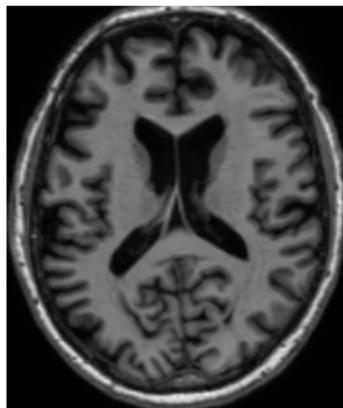


Image size: [494, 424]  
Pixel size: [1.0, 1.0]

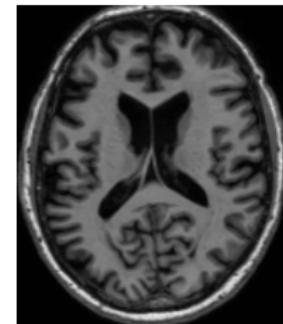


Image size: [395, 339]  
Pixel size: [1.25, 1.25]

- Resample one image to match the *pixel-size* of the other.
- Resize an image with a factor determined by the pixel-size. One example tool in python “scipy.ndimage.zoom”.
- If volumetric images in 3D, then match the *voxel-size*.
- Crop or pad if necessary to make the image sizes the same.

## Exercise

### Images of the same person one year apart

Sizes of the images on the slides are proportional to their number of pixels.



Image size: [454, 384]  
Pixel size: [1.25, 1.25]

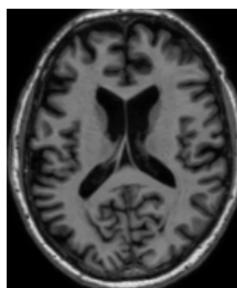


Image size: [494, 424]  
Pixel size: [1.0, 1.0]

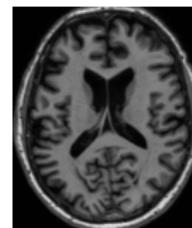


Image size: [395, 339]  
Pixel size: [1.25, 1.25]

- If we are matching the image in the second column to the one in the first one, what are the resize ratios in x- and y-directions?

## Exercise

### Images of the same person one year apart

Sizes of the images on the slides are proportional to their number of pixels.

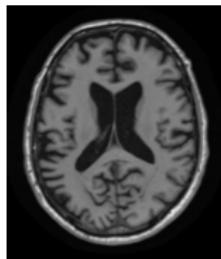


Image size: [454, 384]  
Pixel size: [1.25, 1.25]

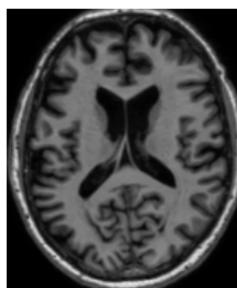


Image size: [494, 424]  
Pixel size: [1.0, 1.0]

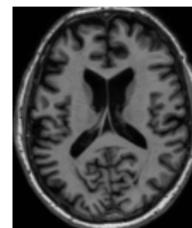


Image size: [395, 339]  
Pixel size: [1.25, 1.25]

- If we are matching the image in the second column to the one in the first one, what are the resize ratios in x- and y-directions? **0.8, 0.8**

## Exercise

### Images of the same person one year apart

Sizes of the images on the slides are proportional to their number of pixels.



Image size: [454, 384]  
Pixel size: [1.25, 1.25]

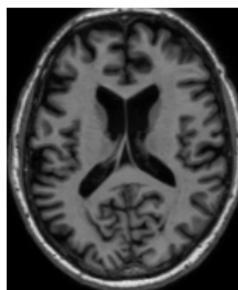


Image size: [494, 424]  
Pixel size: [1.0, 1.0]

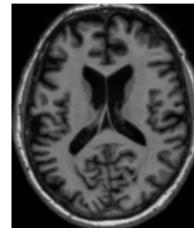


Image size: [395, 339]  
Pixel size: [1.25, 1.25]

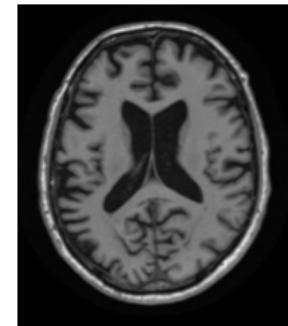


Image size: [568, 480]  
Pixel size: [1.0, 1.0]

- If we are matching the image in the second column to the one in the first one, what are the resize ratios in x- and y-directions? **0.8, 0.8**
- If we are matching the image in the first column to the one in the second one, what are the resize ratios in x- and y-directions?

## Exercise

### Images of the same person one year apart

Sizes of the images on the slides are proportional to their number of pixels.



Image size: [454, 384]  
Pixel size: [1.25, 1.25]

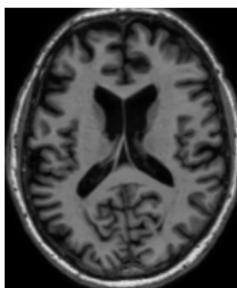


Image size: [494, 424]  
Pixel size: [1.0, 1.0]

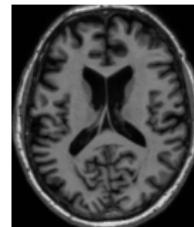


Image size: [395, 339]  
Pixel size: [1.25, 1.25]

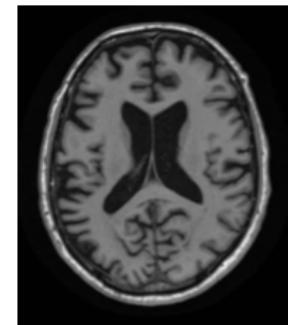


Image size: [568, 480]  
Pixel size: [1.0, 1.0]

- If we are matching the image in the second column to the one in the first one, what are the resize ratios in x- and y-directions? **0.8, 0.8**
- If we are matching the image in the first column to the one in the second one, what are the resize ratios in x- and y-directions? **1.25, 1.25**

## Where do I find this information?

- We need the pixel-size of each image for the matching.
- Such information are stored in the header of the DICOM files.
- It is also stored in the headers of many volumetric medical image formats.
- One way to reach them is to use the “nibabel” package in python.

## How **not** to tackle it.

**Images of the same person one year apart**

Sizes of the images on the slides are proportional to their number of pixels.



Image size: [454, 384]

Pixel size: [1.25, 1.25]

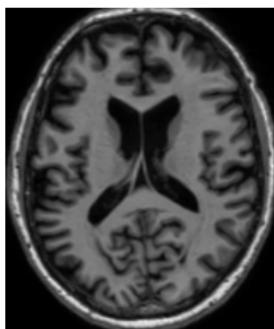


Image size: [494, 424]

Pixel size: [1.0, 1.0]

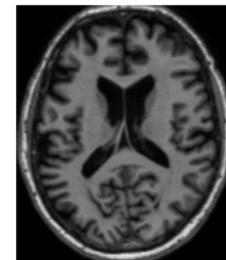


Image size: [395, 339]

Pixel size: [1.25, 1.25]

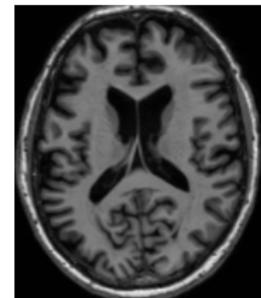


Image size: [454, 384]

Pixel size: [1.09, 1.11]

- A very naive way of doing the matching is through matching the image sizes.
- Last column on the right is obtained by matching the size of the image on the second column to that in the first one.
- Notice that the images are not comparable any more.
- Even the aspect ratio changed.

## How not to tackle it.

### Images of the same person one year apart

Sizes of the images on the slides are proportional to their number of pixels.



Image size: [454, 384]  
Pixel size: [1.25, 1.25]

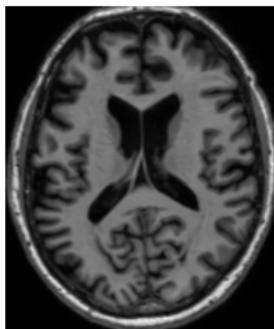


Image size: [494, 424]  
Pixel size: [1.0, 1.0]

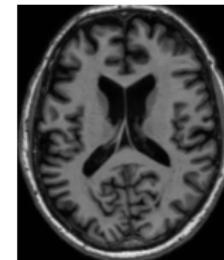


Image size: [395, 339]  
Pixel size: [1.25, 1.25]

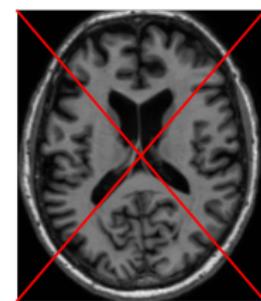


Image size: [454, 384]  
Pixel size: [1.09, 1.11]

- A very naive way of doing the matching is through matching the image sizes.
- Last column on the right is obtained by matching the size of the image on the second column to that in the first one.
- Notice that the images are not comparable any more.
- Even the aspect ratio changed.

### Important

Always match the pixel / voxel-size. Do not match the image size, it may lead to wrong normalization.