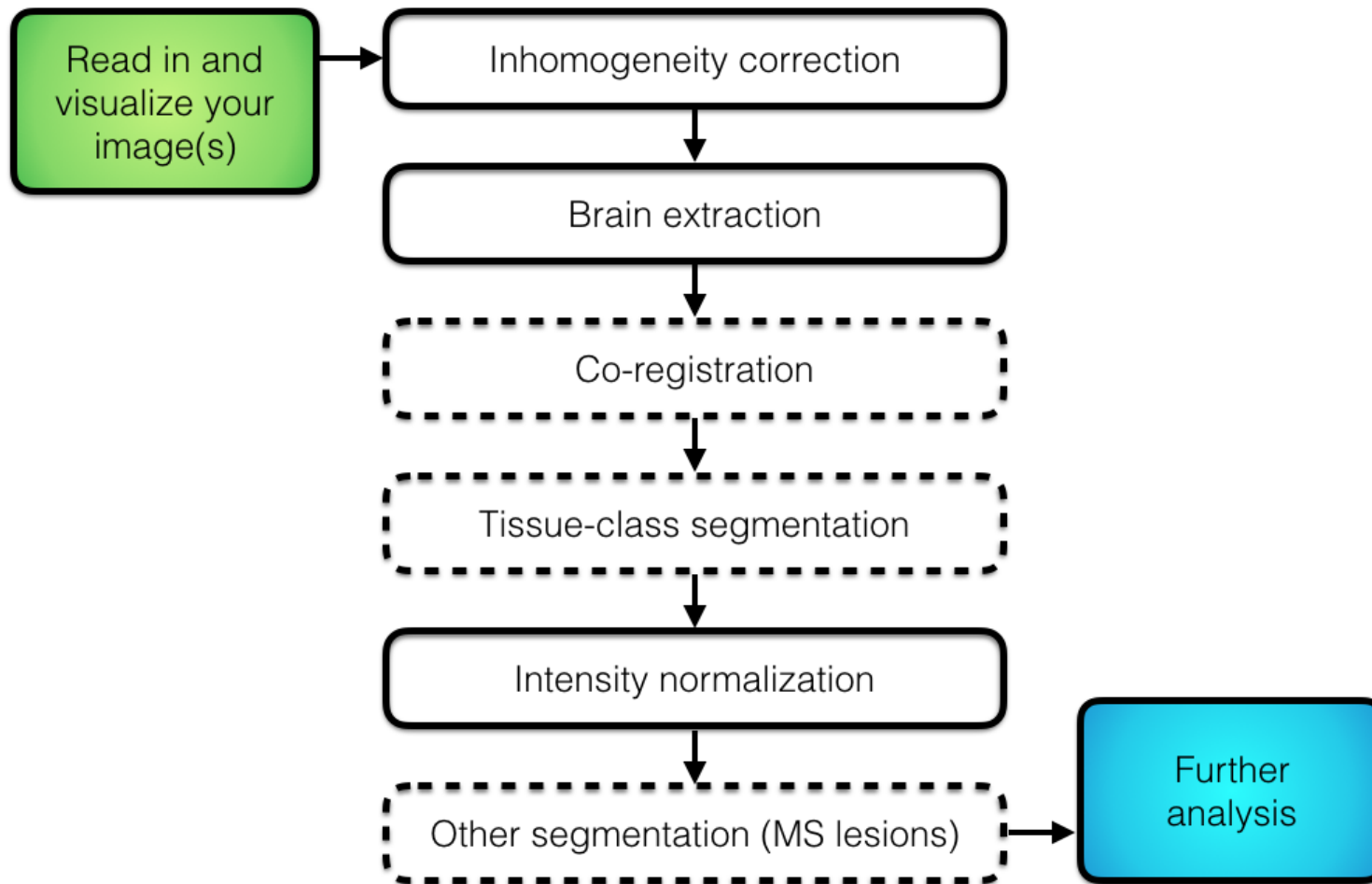# Tissue-Class Segmentation

Overall Pipeline

Image segmentation

- We are often interested in subdividing or segmenting the brain into meaningful biological regions of interest (ROIs) for an analysis.

- Examples: tissue segmentation, segmentation of gray matter structures, segmentation of pathology (MS lesions, tumors, …)

- We will perform 3-class tissue segmentation in R using `fslr` and `ANTsR`:

    - Cerebrospinal fluid (CSF)

    - Gray Matter (GM)

    - White Matter (WM)

Loading Data

· Let's read in the training T1 and brain mask for subject 05 (not 01!).

```r
library(ms.lesion)
library(neurobase)
all_files = get_image_filenames_list_by_subject(
  group = "training",
  type = "coregistered")
files = all_files$training05 # NOT training subject 1!
t1 = readnii(files["T1"])
rt1 = robust_window(t1)
mask = readnii(files["Brain_Mask"])
```

```r
run_mask = t1 > 100
dd_orig = drop_empty_dim(run_mask, keep_ind = TRUE)
```
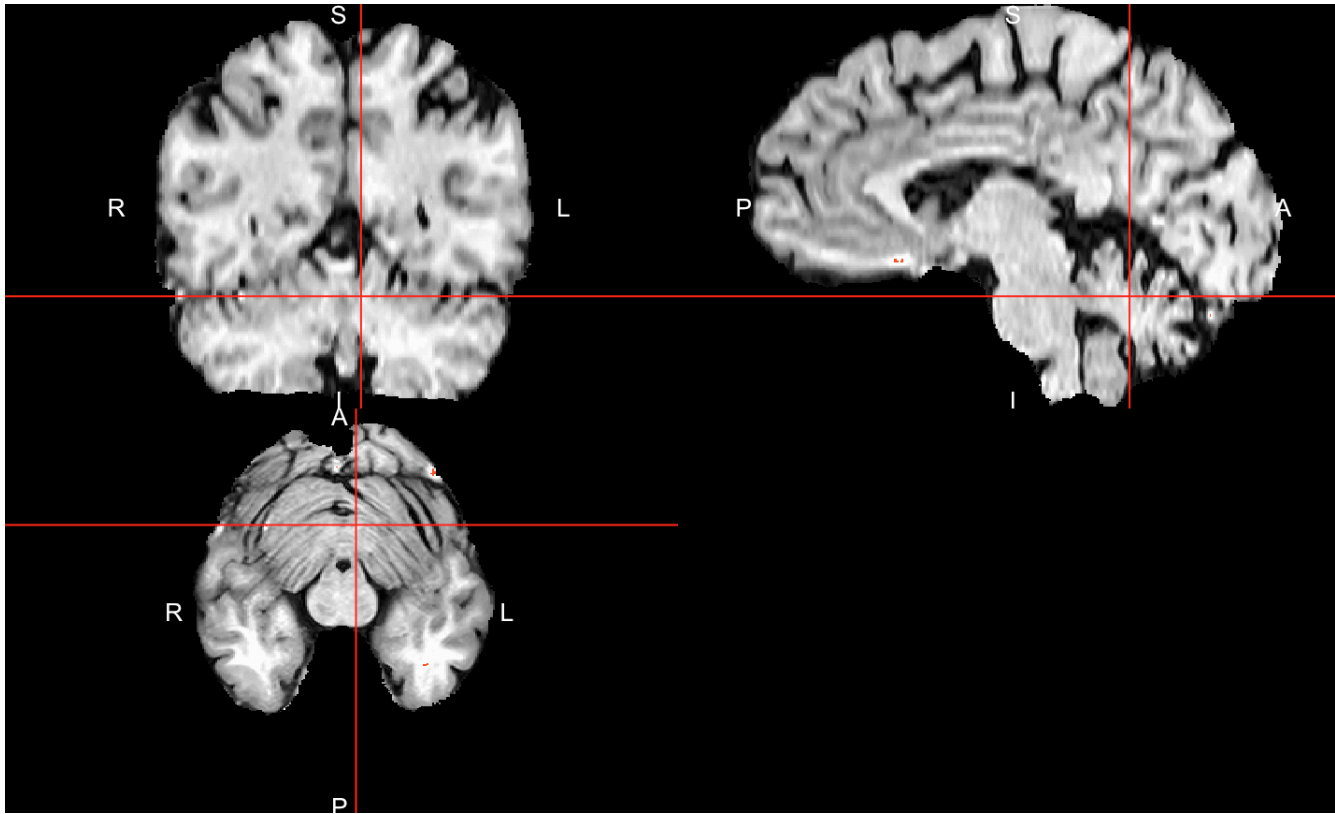
Tissue Segmentation: Large Outliers

- Many tissue class segmentations are based on k-means clustering.
- These methods can be skewed by large outliers.

```
hist(t1, mask = mask, breaks = 2000); text(x = 600, y = 3000, '"outliers"')
```
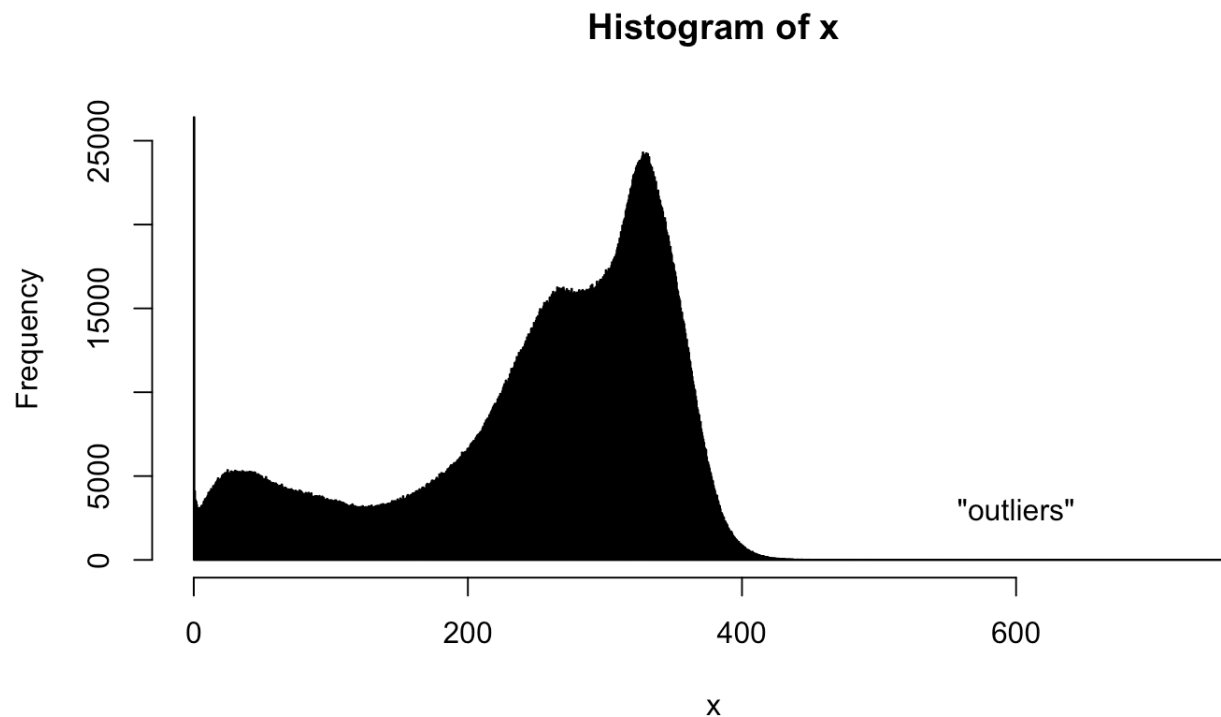

Histogram of x

Where are the outliers?

```
ortho2(rt1, t1 > 450, xyz = xyz(t1 > 450)) # xyz - cog of a region
```
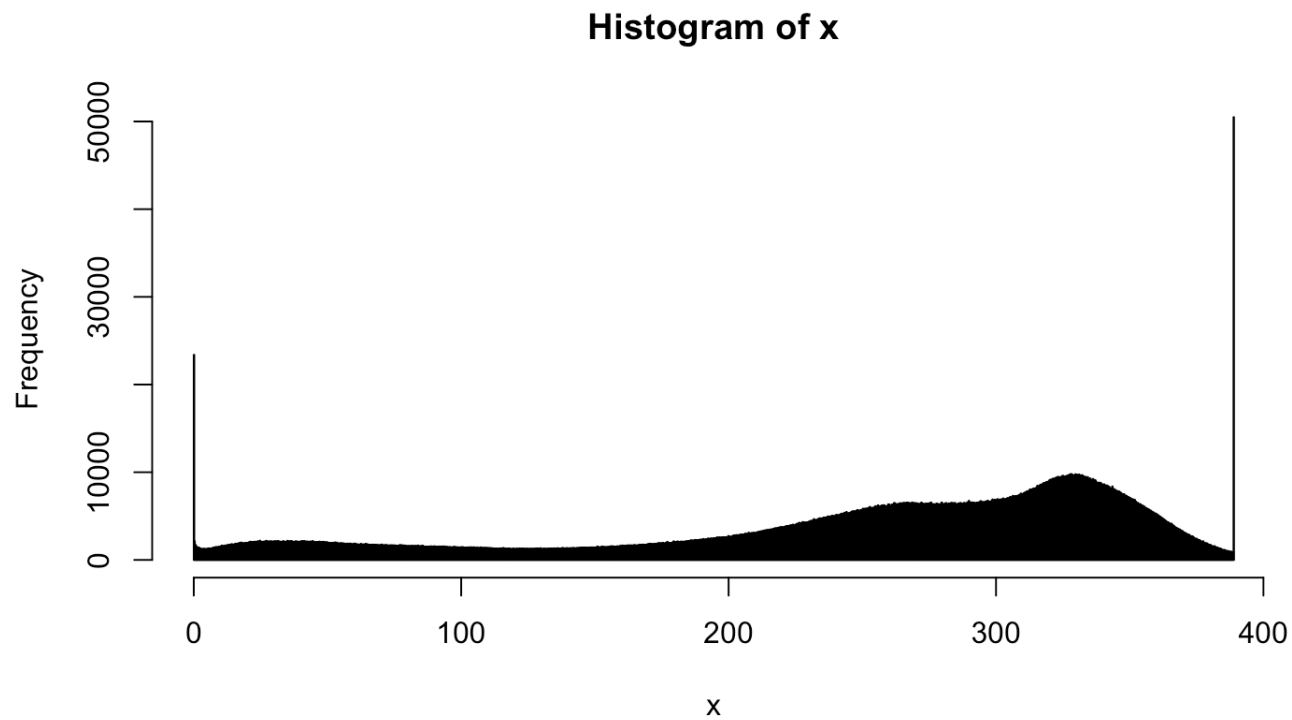
Cleaning up values: Masking the image

```
t1[ t1 < 0 ] = 0
t1 = mask_img(t1, mask)
rt1 = robust_window(t1)
```

What does the histogram look like now?

```
hist(rt1, mask = mask, breaks = 2000);
```

Histogram of x

Tissue Segmentation using FSL FAST

- FAST is based on a hidden Markov random field model and an Expectation-Maximization algorithm (Zhang, Brady, and Smith 2001).

- It jointly produces a bias field corrected image and a probabilistic tissue segmentation.

- More robust to noise and outliers than finite mixture model-based methods that do not incorporate spatial information.

The `fslr` function `fast` calls fast from FSL. The `--nobias` option tells FSL to not perform inhomogeneity correction (N4 already performed in `ANTsR`).

```
t1file = files["T1"]
t1fast = fast(t1,
          outfile = paste0(nii.stub(t1file), "_FAST"),
          opts = "--nobias")
```
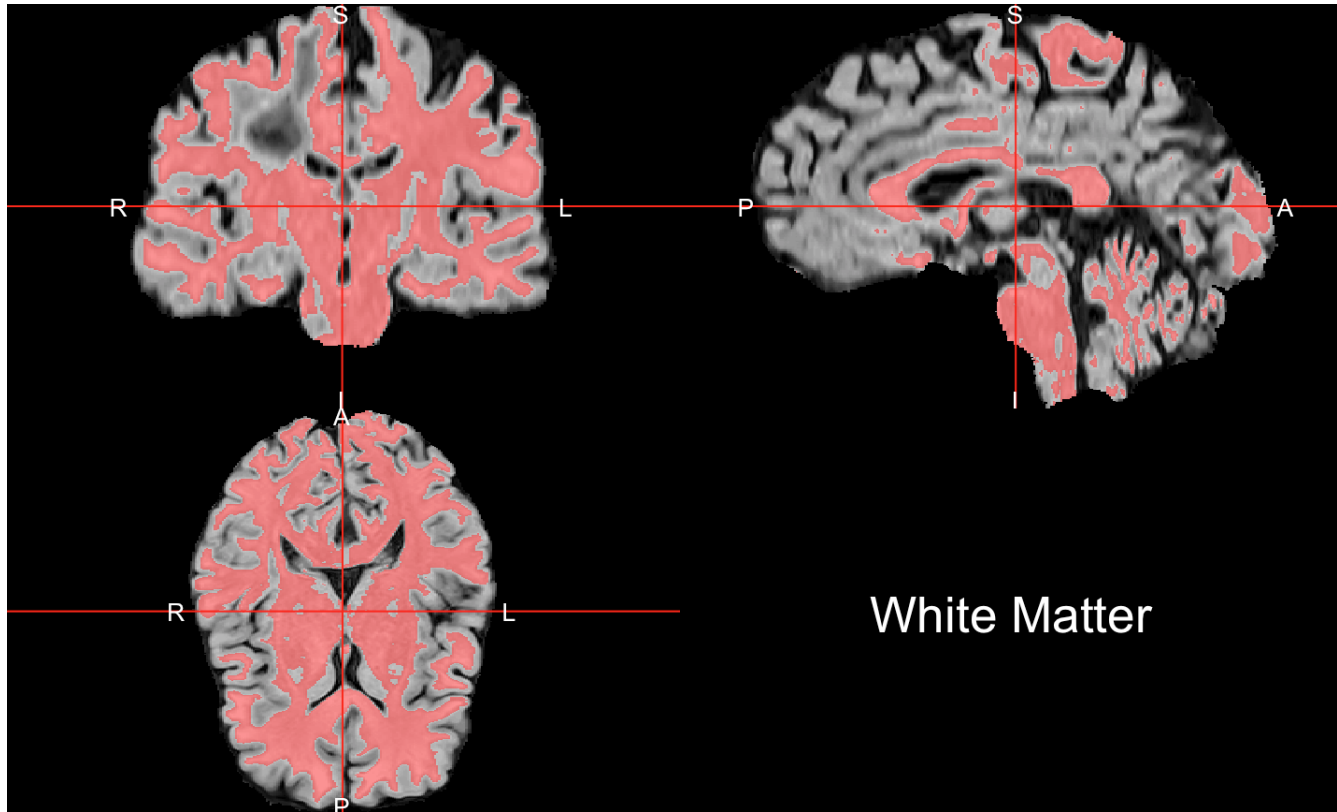
FAST Results

FAST assumes three tissue classes and produces an image with the three labels, ordered by increasing within-class mean intensities. In a T1 image, this results in:

- Level 1: CSF
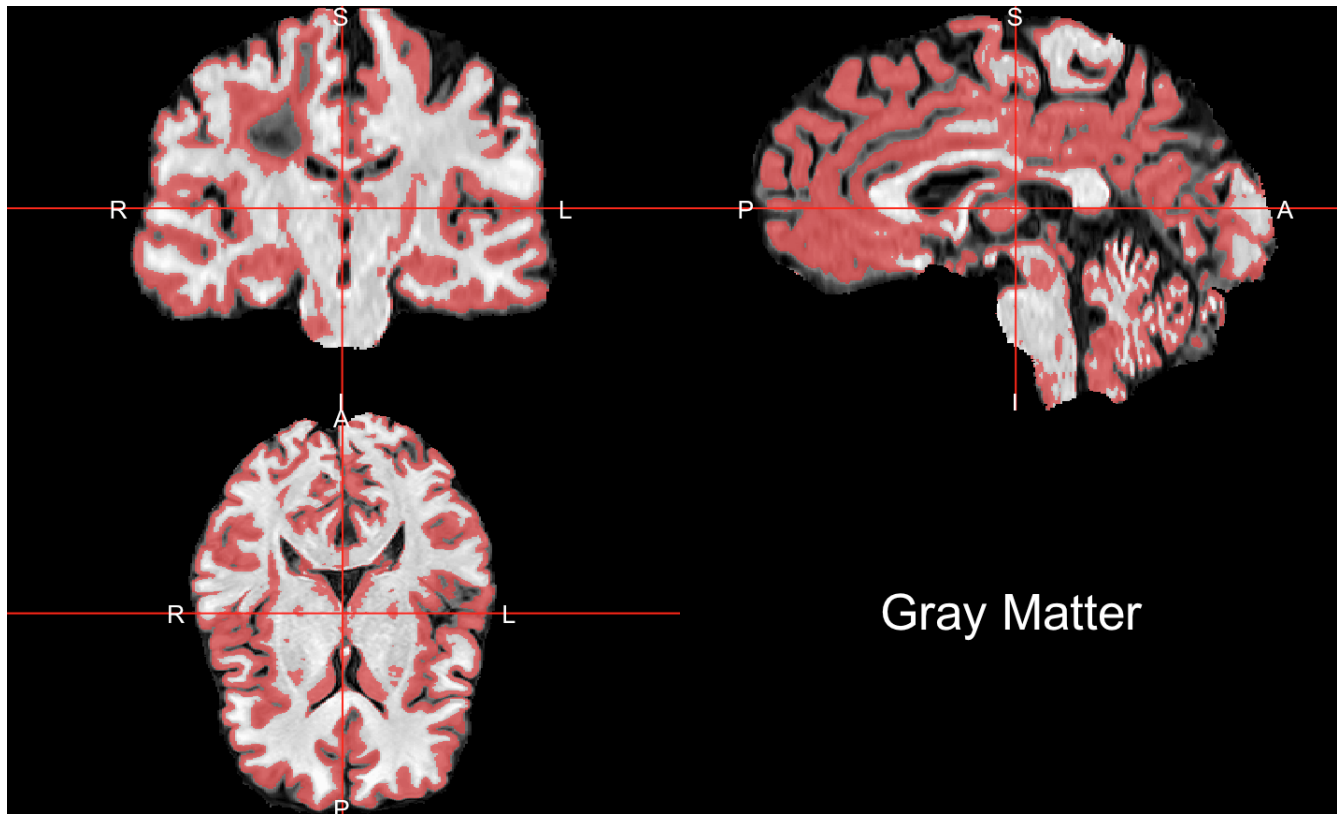- Level 2: Gray Matter
- Level 3: White Matter

FAST: White Matter

```
ortho2(rt1, t1fast == 3, col.y = alpha("red", 0.5), text = "White Matter")
```
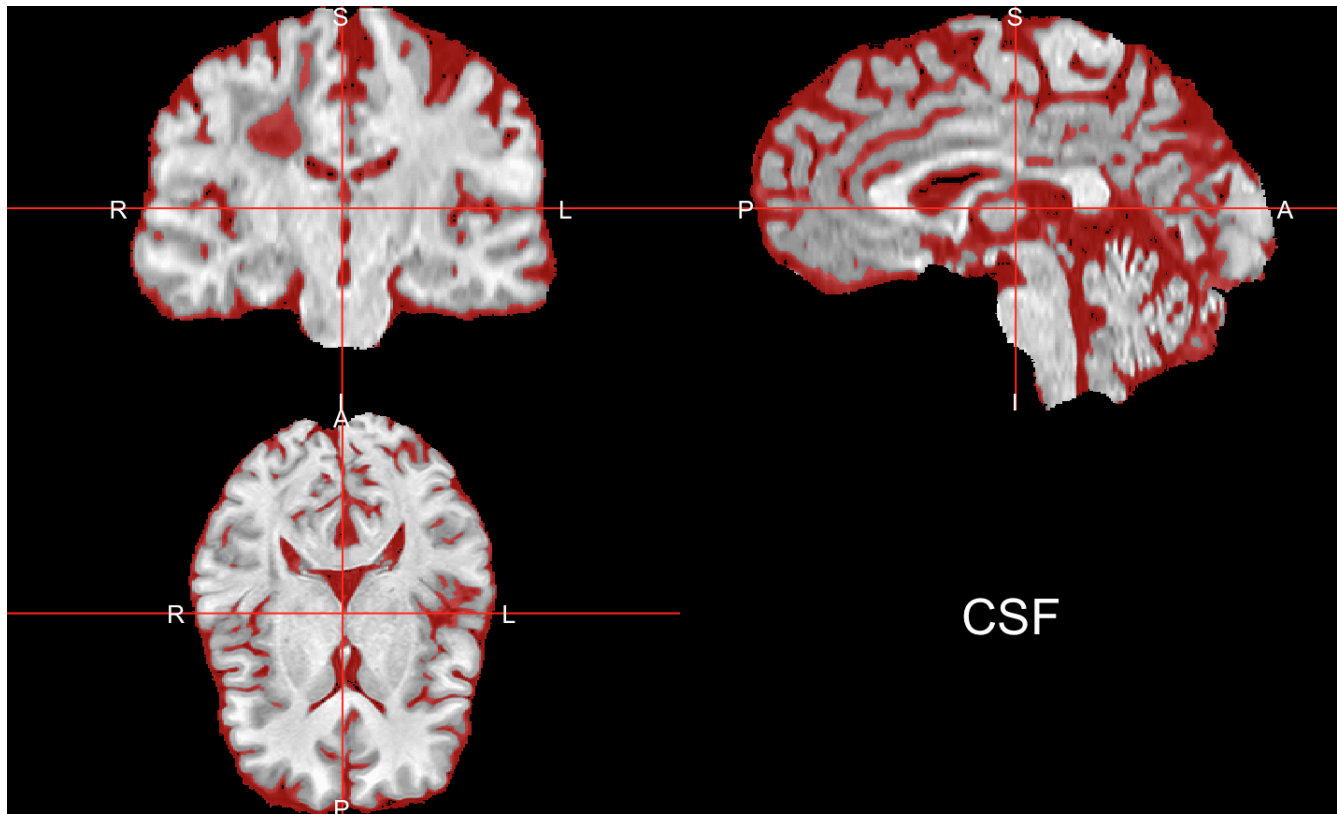
FAST: Gray Matter

```r
ortho2(rt1, t1fast == 2, col.y = alpha("red", 0.5), text = "Gray Matter")
```
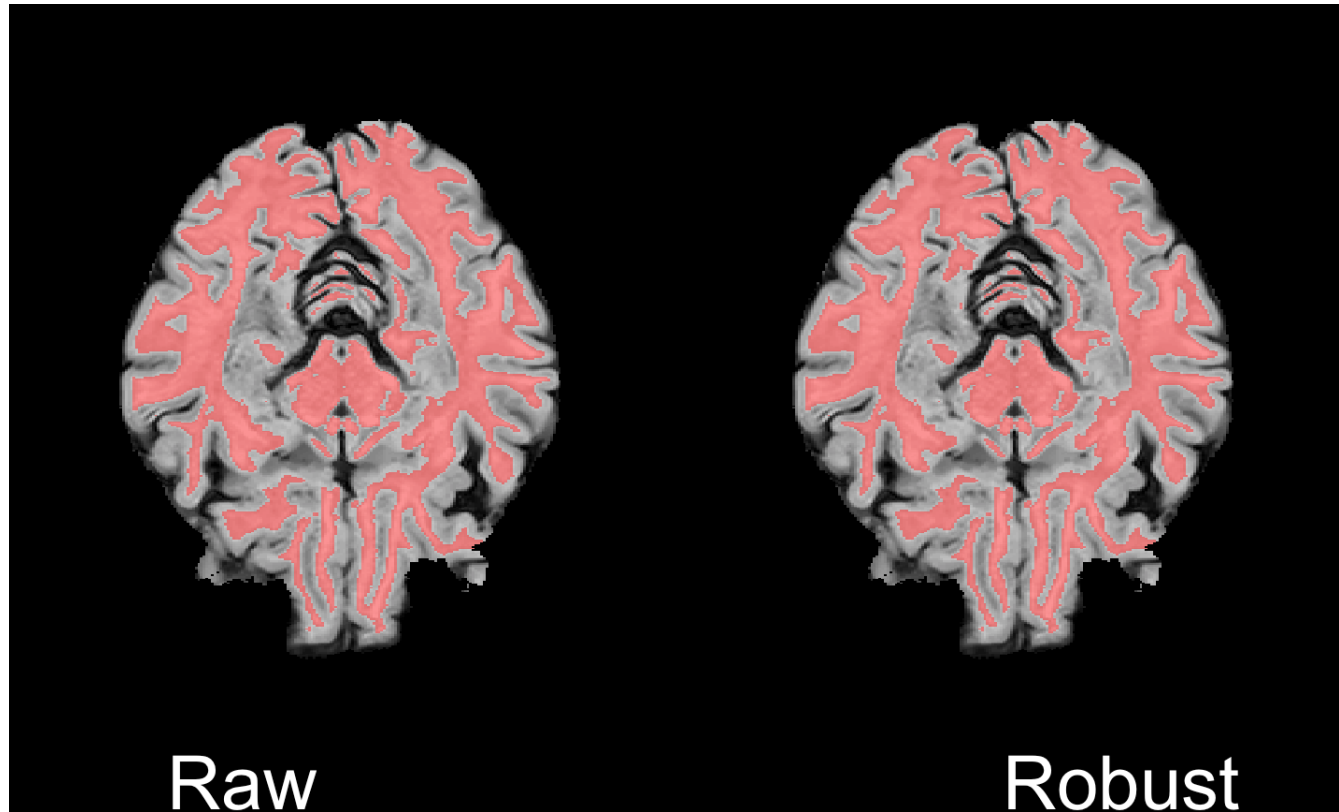
FAST: CSF

```
ortho2(rt1, t1fast == 1, col.y = alpha("red", 0.5), text = "CSF")
```
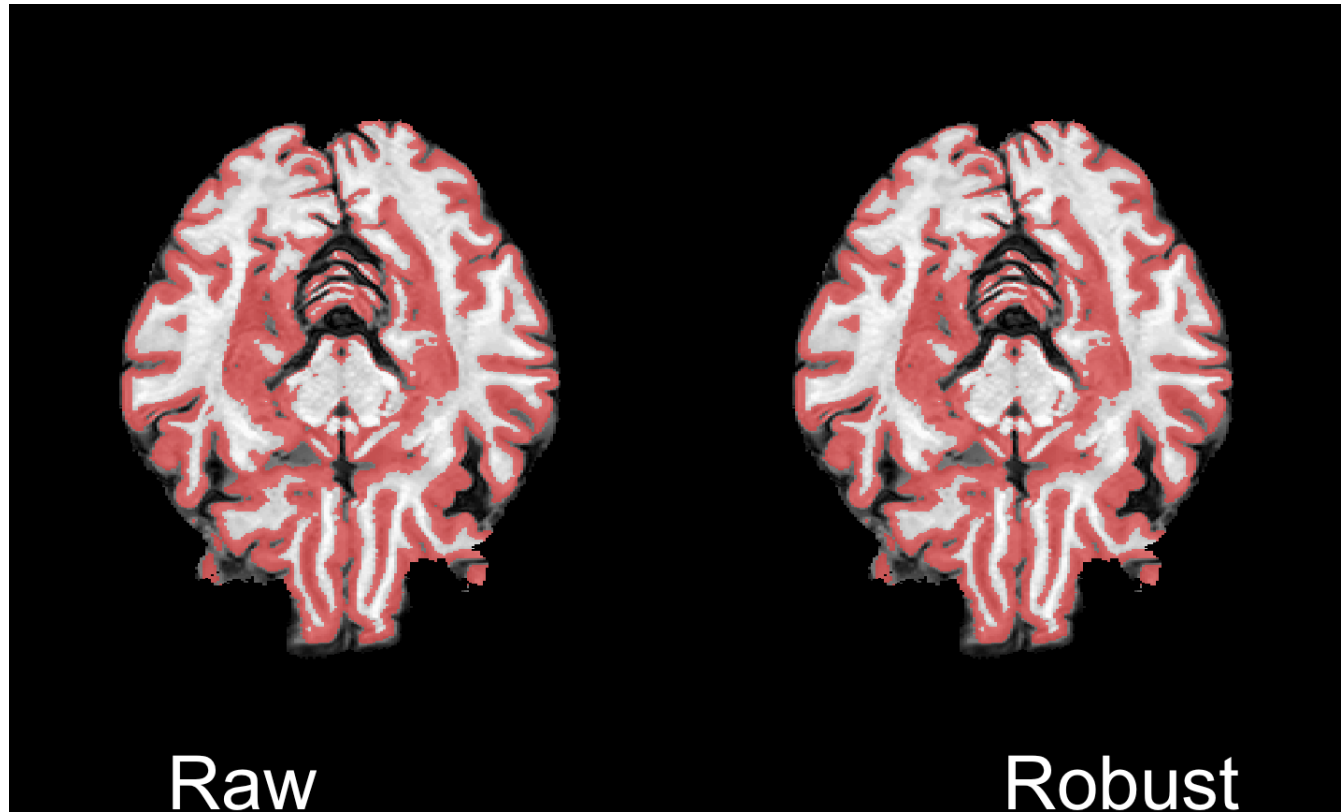
Removing large values: Is there an effect?

```
robust_fast = fast(rt1, # the robust_window(t1)
                   outfile = paste0(nii.stub(t1file), "_FAST"),
                   opts = "--nobias")
```
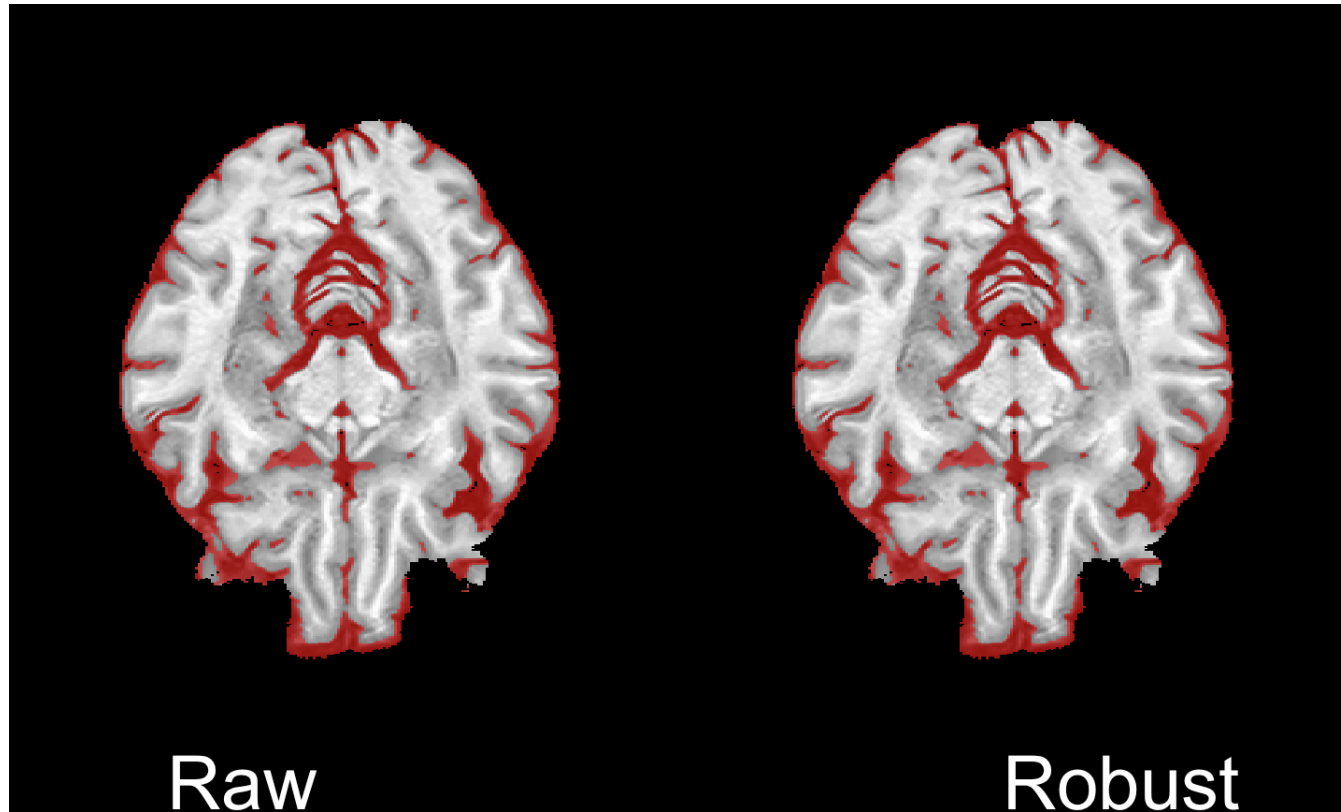
FAST with Window: White Matter

Raw

Robust

FAST with Window: Gray Matter

FAST with Window: CSF

FAST Results

- Overall the results look good
 - Not much difference after dampening outliers using `robust_window`

Tissue Segmentation using ANTsR, extrantsr

- Uses Atropos (Avants et al. 2011)
    - 3D K-means clustering + a Markov random field
- The `extrantsr::otropos` function works with `nifti` objects
    - calls `ANTsR::atropos` function

```r
t1_otropos = otropos(a = t1, x = mask) # using original data
t1seg = t1_otropos$segmentation
```
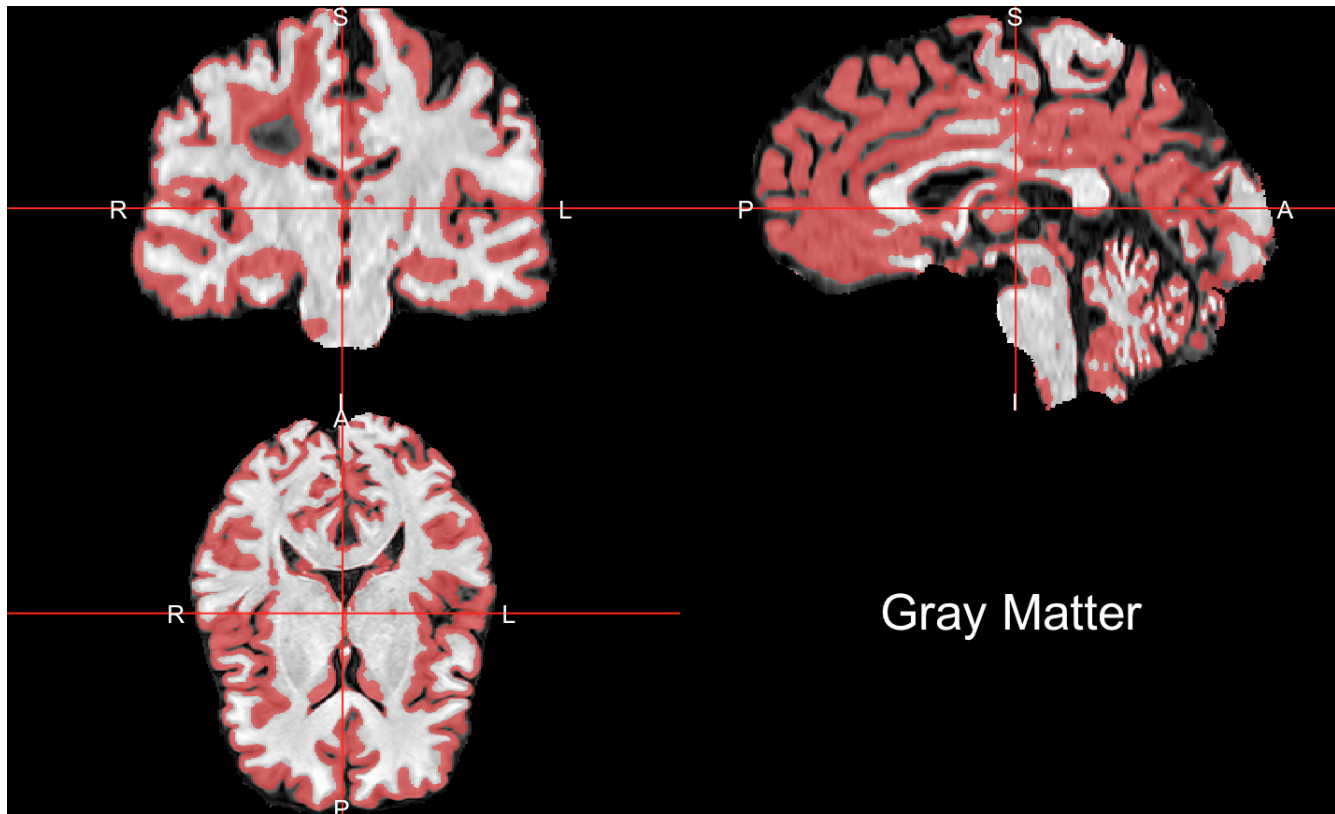
Atropos: White Matter

```
ortho2(rt1, t1seg == 3, col.y = alpha("red", 0.5), text = "White Matter")
```
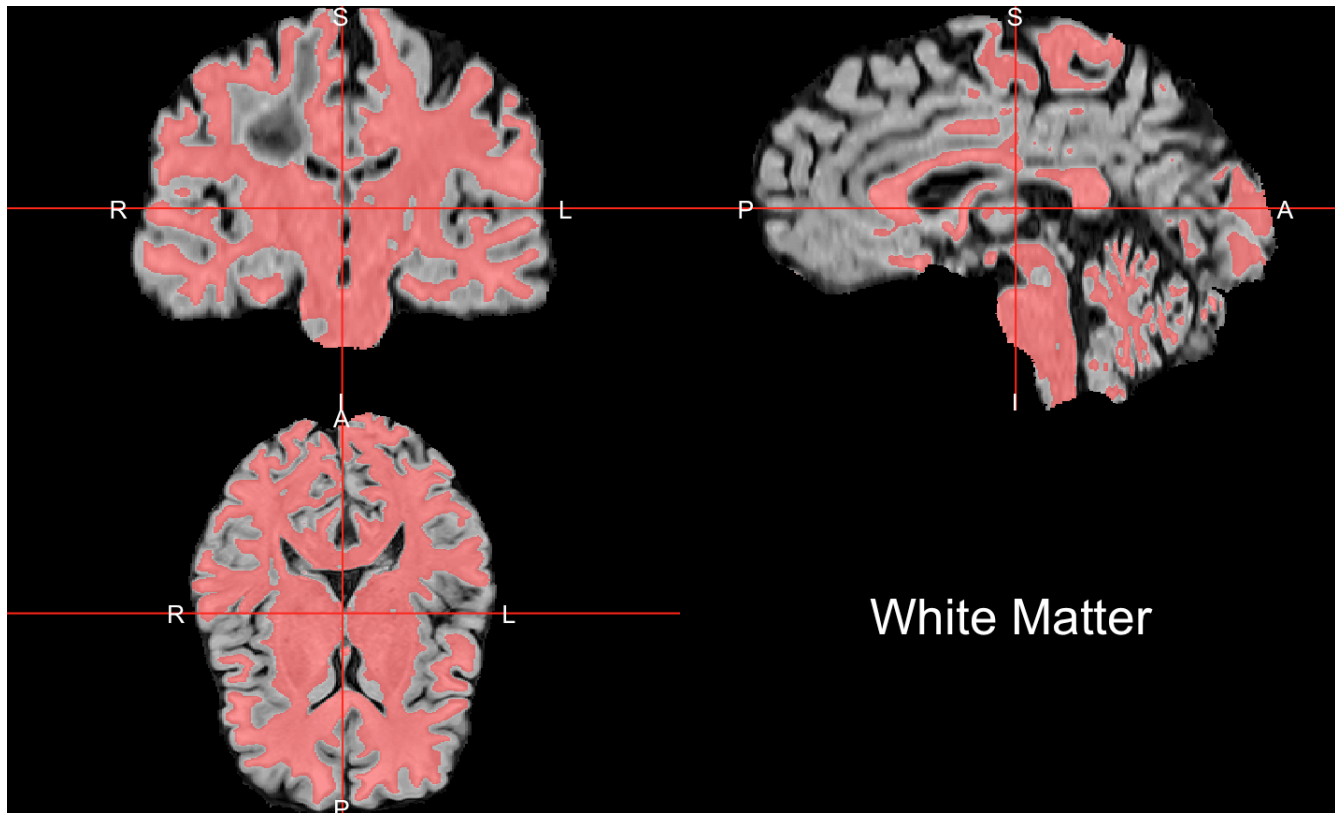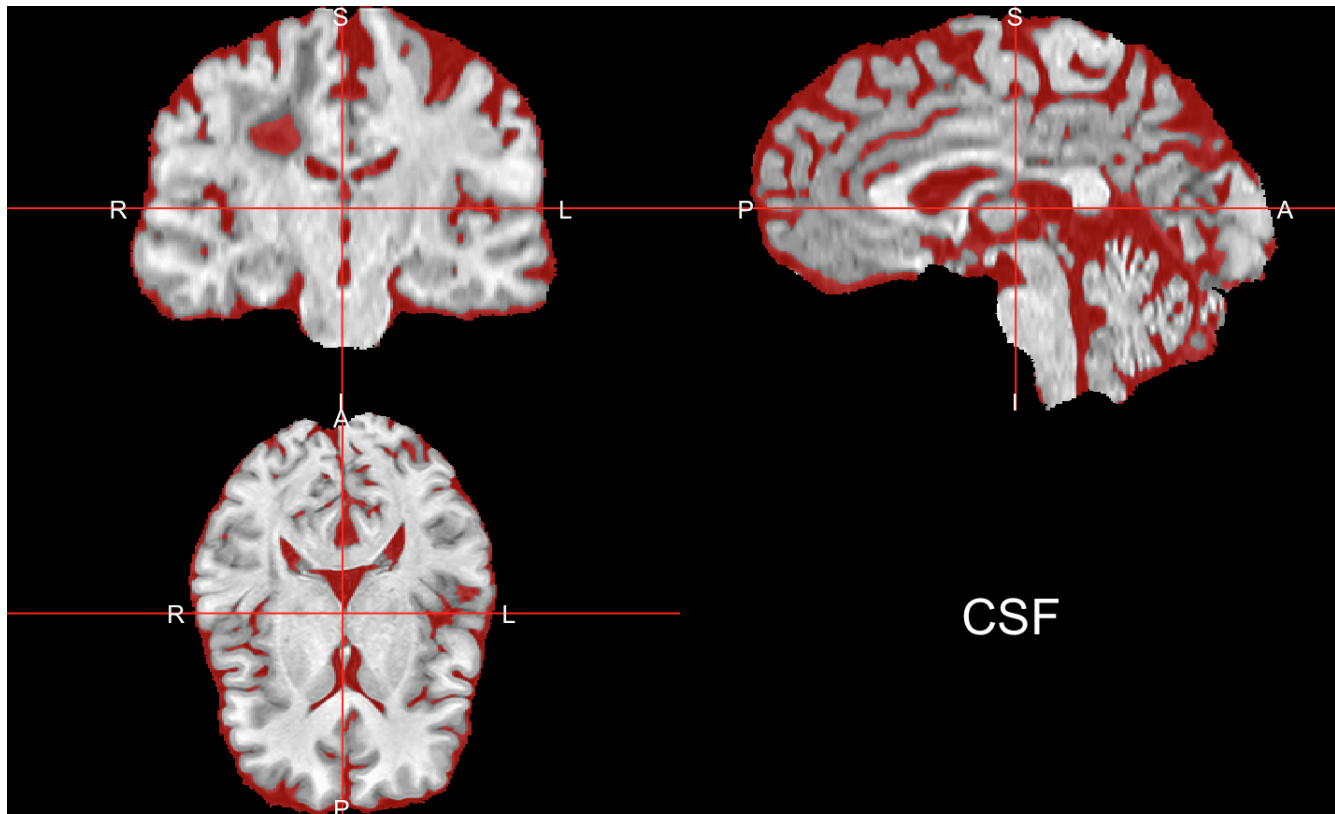
Atropos: Gray Matter

```
ortho2(rt1, t1seg == 2, col.y = alpha("red", 0.5), text = "Gray Matter")
```

Atropos: CSF

```
ortho2(rt1, t1seg == 1, col.y = alpha("red", 0.5), text = "CSF")
```
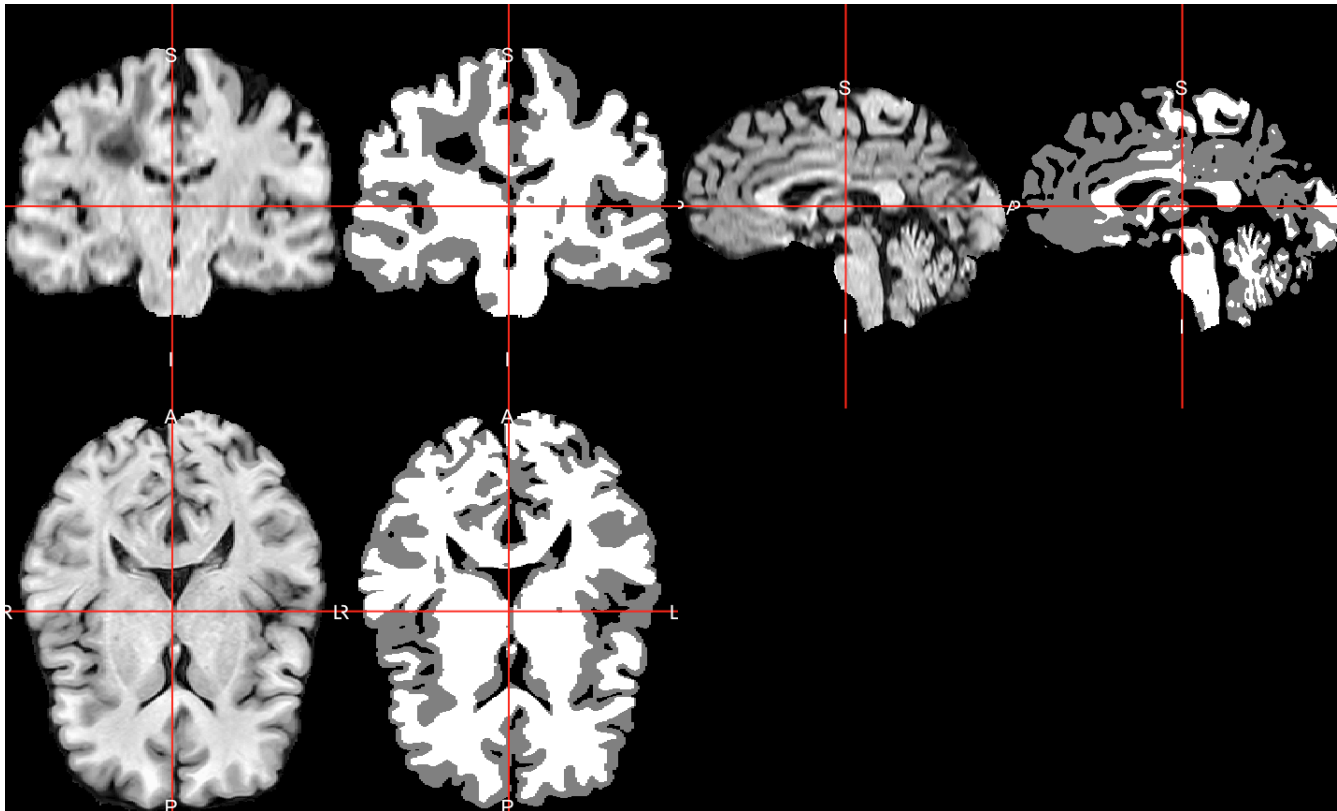
Default Atropos Results

- Overall the results do not look good
    - The k-means clustering is affected by large outliers
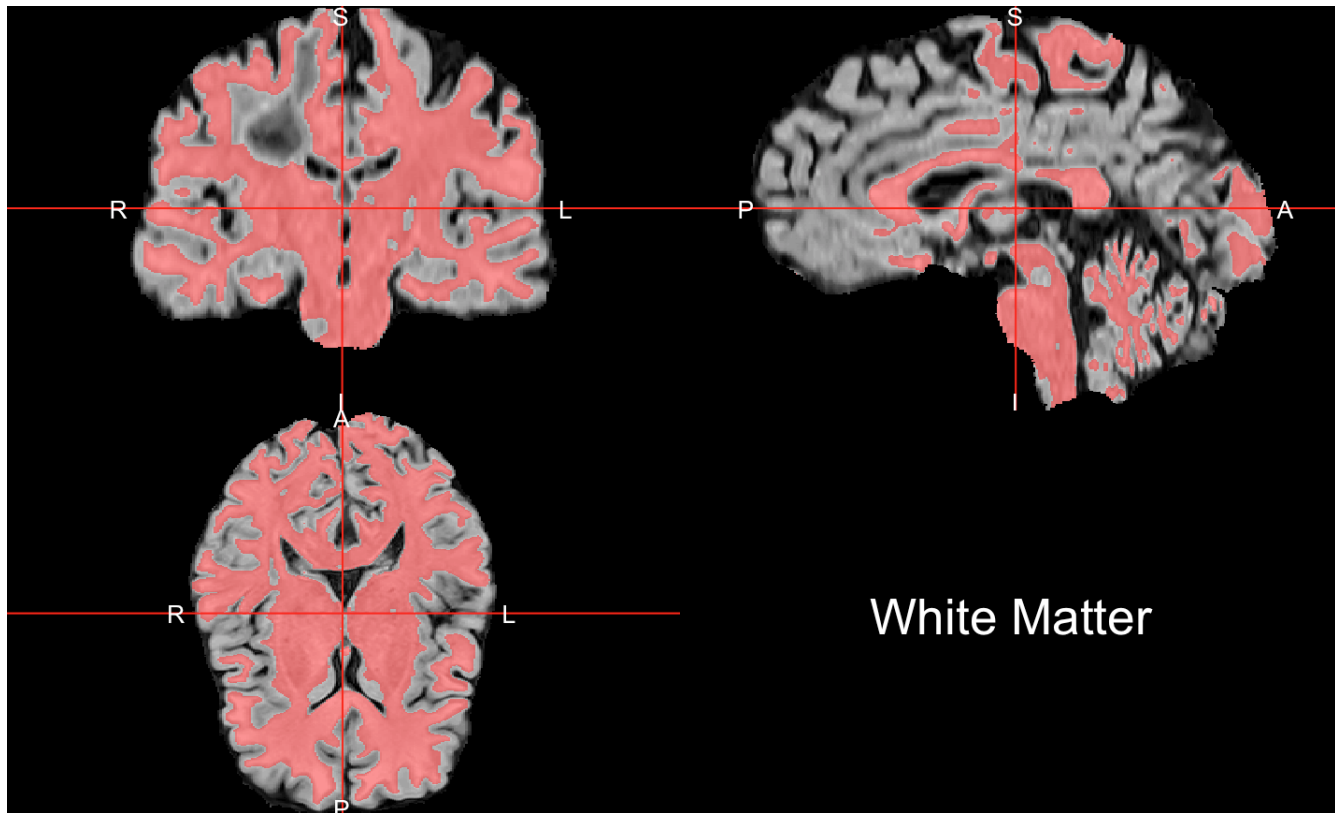- We will try using `robust_window`

Atropos using Windowing

```
robust_t1_otropos = otropos(a = rt1, x = mask) # using robust
robust_t1seg = robust_t1_otropos$segmentation
```

```
double_ortho(rt1, robust_t1seg)
```
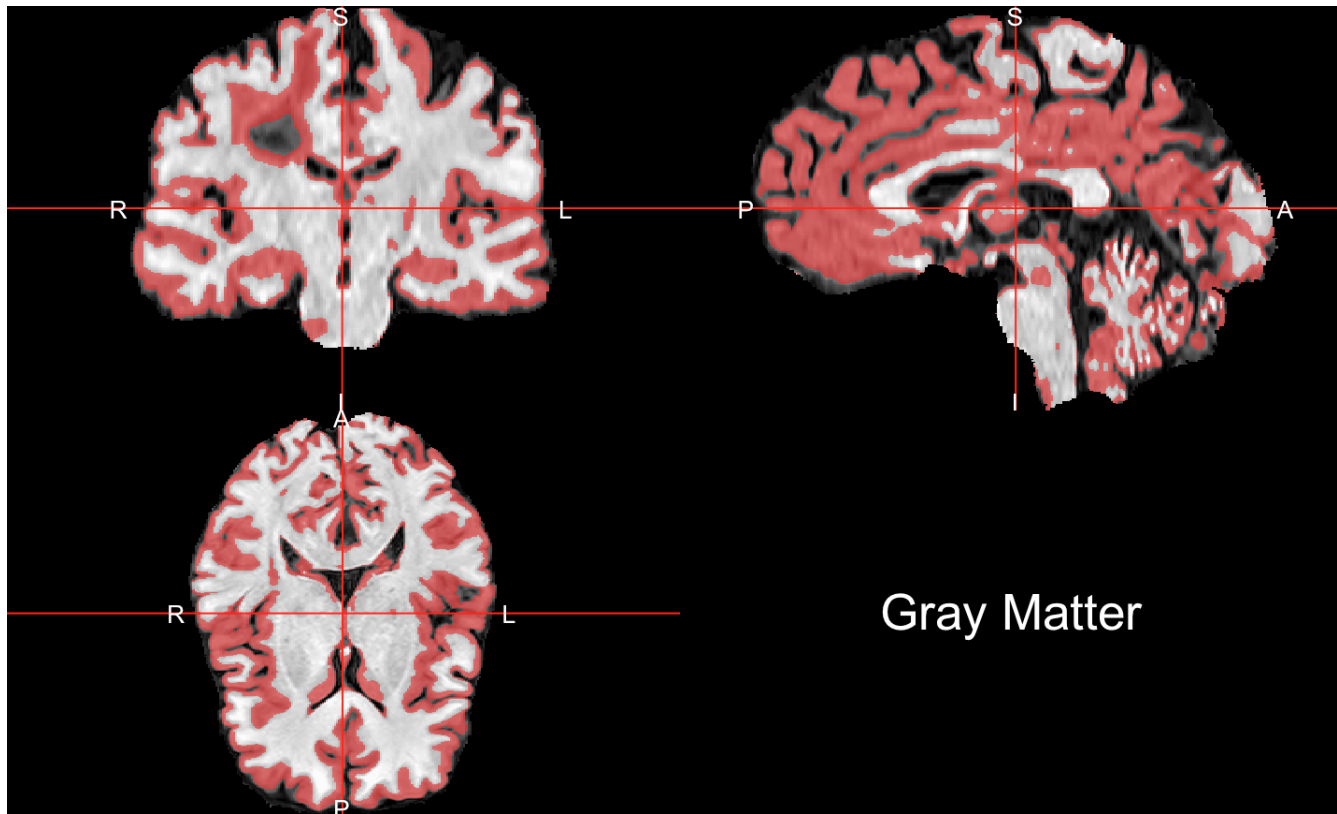
Atropos with Window: White Matter

```
ortho2(rt1, robust_t1seg == 3, col.y = alpha("red", 0.5), text = "White Matter")
```
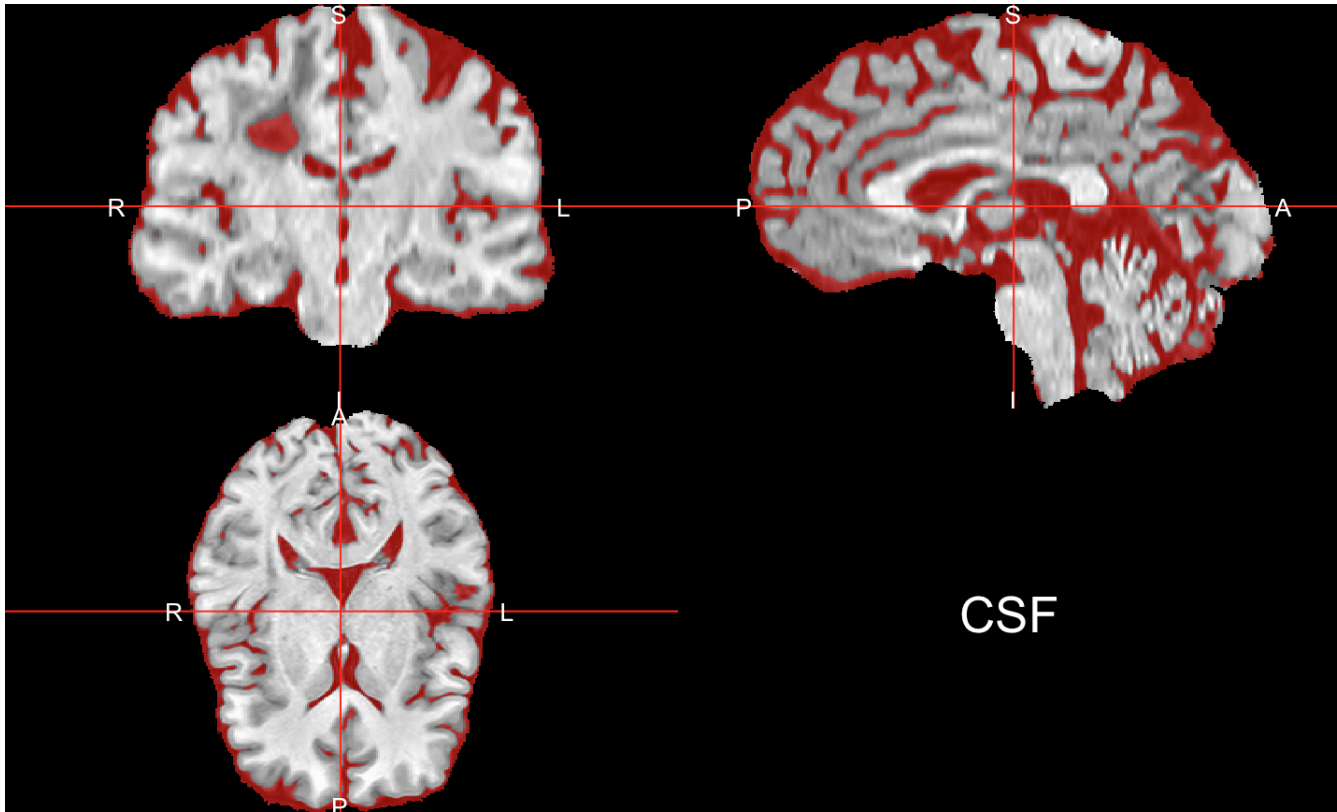
Atropos with Window: Gray Matter

```
ortho2(rt1, robust_t1seg == 2, col.y = alpha("red", 0.5), text = "Gray Matter")
```

Atropos with Window: CSF
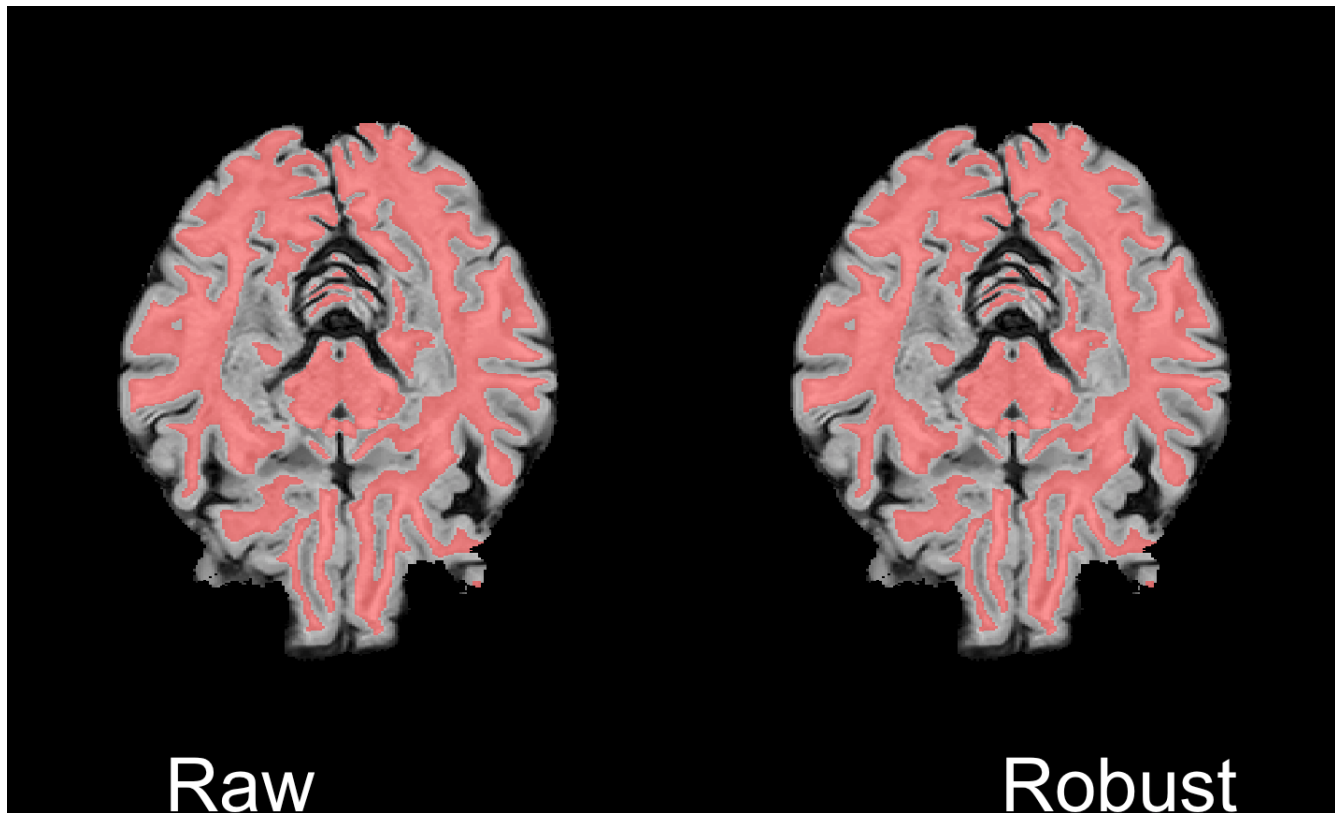
```r
ortho2(rt1, robust_t1seg == 1, col.y = alpha("red", 0.5), text = "CSF")
```
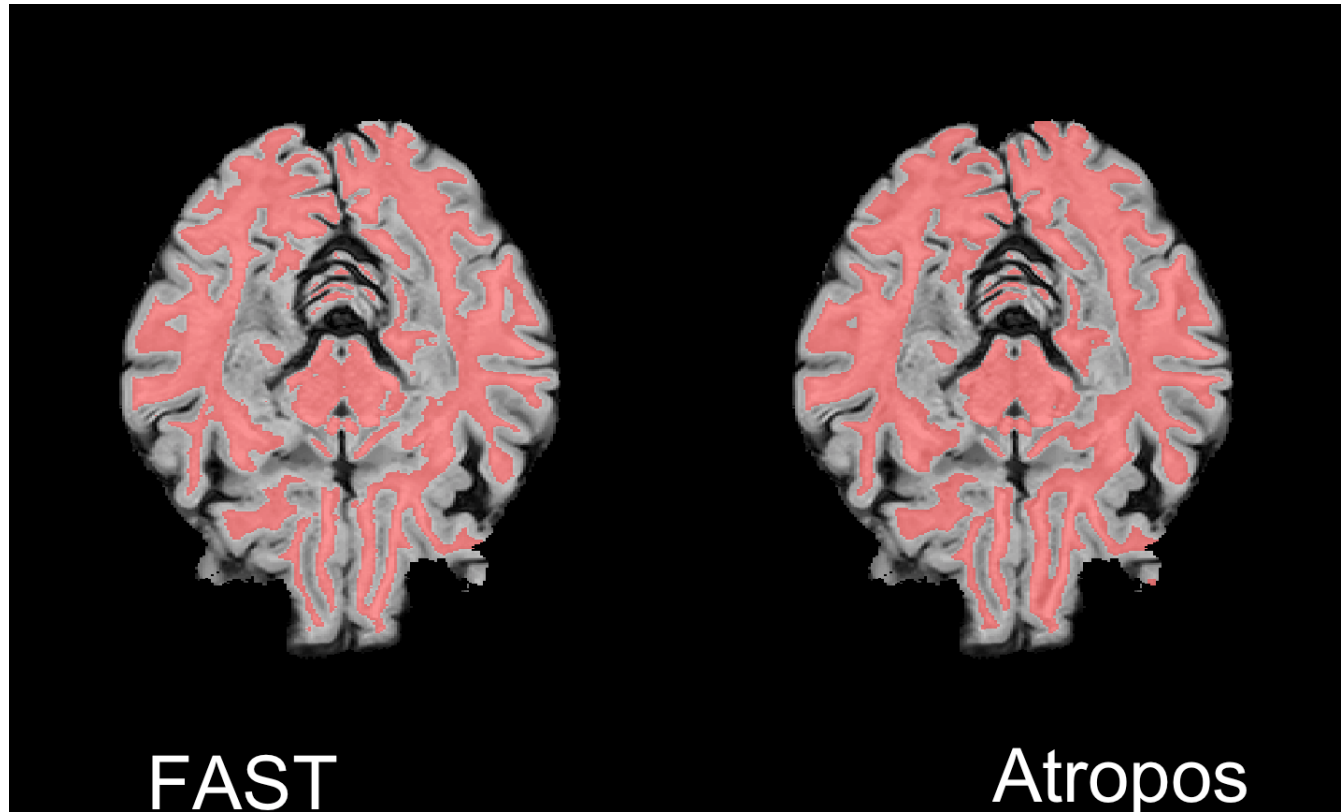
Atropos with Window Results

- Overall the results look like they reasonably separate the classes
    - No ground truth
- Winsorizing large outliers aided the k-means clustering
    - Results much better than running Atropos on the raw data

Raw   Robust

Atropos WM vs. FAST WM

FAST

Atropos

Estimating the Volume of Each Class

We can create a table which will count the number of voxels in each category:

```
tab_fsl = table(robust_fast[ robust_fast != 0])
tab_ants = table(robust_t1seg[ robust_t1seg != 0])
names(tab_fsl) = names(tab_ants) = c("CSF", "GM", "WM")
tab_fsl
```

```
    CSF      GM      WM
1551816 2632937 2932684
```

```
tab_ants
```

```
    CSF      GM      WM
1288931 2545481 3304745
```

```
out_mask = robust_fast != 0 | robust_t1seg != 0
table(robust_fast[ out_mask ], robust_t1seg[ out_mask ])
```

```
         1        2        3
0    21720        0        0
1  1267209   284607        0
2        2  2257284   375651
3        0     3590  2929094
```

Estimating the Volume of Each Class

By multiplying by the voxel resolution (in cubic centimeters) using the `voxres` function, we can get volumes

```
vres = oro.nifti::voxres(t1, units = "cm")
print(vres)
```

```
[1] 0.0001757813
```

```
vol_fsl = tab_fsl * vres
vol_fsl
```

```
     CSF        GM        WM
272.7802  462.8210  515.5109
```

```
vol_ants = tab_ants * vres
vol_ants
```

```
     CSF        GM        WM
226.5699  447.4478  580.9122
```

Website

http://johnmuschelli.com/imaging_in_r

References

Avants, Brian B, Nicholas J Tustison, Jue Wu, Philip A Cook, and James C Gee. 2011. "An Open Source Multivariate Framework for N-Tissue Segmentation with Evaluation on Public Data." 9 (4). Springer:381–400.

Zhang, Yongyue, Michael Brady, and Stephen Smith. 2001. "Segmentation of Brain MR Images Through a Hidden Markov Random Field Model and the Expectation-Maximization Algorithm." 20 (1):45–57. http://ieeexplore.ieee.org/xpls/abs_all.jsp?arnumber=906424.