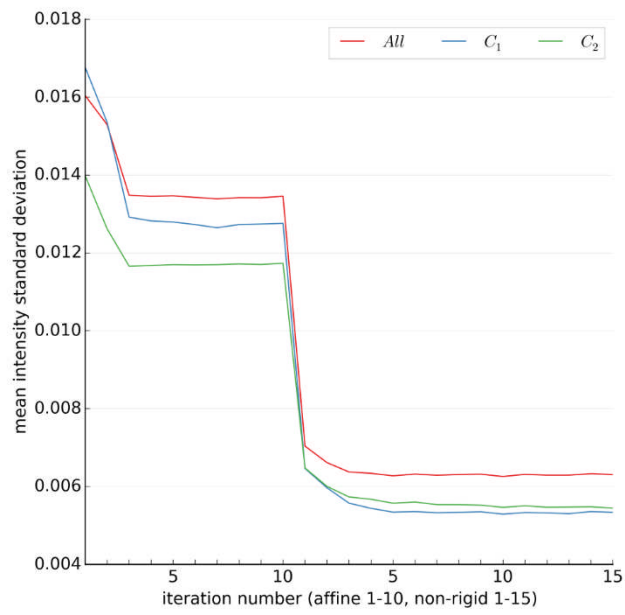


## 1 **S1. Group-wise registration assessment**

2 We quantified the progressive improvement in image alignment at each iteration of group-wise  
3 registration (GWR), and justified the number of iterations, by measuring the intensity standard  
4 deviation  $\sigma_I$  between resampled images at every voxel within the brain. Inter-image intensities in  
5 equivalent aligned regions should have low standard deviation, thanks to standardisation prior to  
6 GWR. Fig A shows the mean  $\sigma_I$  over all voxels, at each iteration, within the brain mask, for all brains,  
7 and for  $C_1$  and  $C_2$ . After a dramatic decrease with the first iteration of NRR,  $\sigma_I$  reaches a plateau after  
8 5-10 iterations.

9 We sought to assess whether the difference in SNR and CNR between cohorts affected the precision  
10 of spatial alignment. We resampled each binarised brain mask into the final GWR average space. For  
11 each resampled mask, we computed the Jaccard index with every other resampled mask in the cohort,  
12 and calculated their mean and standard deviation. We compared cohorts' mean Jaccard indices with  
13 a t-test. There was no appreciable difference ( $p > 0.9$ ; Cohen's  $d = 0.0065$ ): GWR performed similarly in  
14 aligning brain masks from each cohort.

15 We also used the per-voxel standard deviation to compare cohorts' alignment (Fig A). Deviation in  
16 voxel intensities should arise from alignment offset, natural variations in local tissue intensity, and  
17 noise. Fig A illustrates that after NRR,  $C_1$  and  $C_2$  had similar mean standard deviations, over the whole  
18 brain.



19

20 Fig A: **Group-wise registration assessment:** Mean standard deviation of brain voxels in resampled  
 21 images, within the brain mask, for all brains and each cohort individually.