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Improving skullstripping and nonlinear warping in AFNI: sswarper2

National Institute of Mental Health

Results

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Here we describe *sswarper2*, a new nonlinear alignment program in AFNI¹ that both skullstrips (SS) a T1w volume and aligns (warps) it to a reference dataset. We show its improvements to predecessor @*SSwarper*, which had similar dual roles.

Methods & Results

Introduction

The research and writing of the manuscript were supported by the NIMH IRP (ZICMH002888) of the NIH (HHS, USA). This work utilized the computational resources of the NIH HPC Biowulf cluster ([http://hpc.nih.gov\)](http://hpc.nih.gov/). [1] Cox RW (1996). Comput Biomed Res 29(3):162-173. [2] Cox RW, Glen DR (2013). Proc. OHBM-2013. [3] Saad ZS, Glen DR, Chen G, Beauchamp MS, Desai R, Cox RW (2009). Neuroimage 44 839–848. [4] Taylor PA, Glen DR, Reynolds RC, Basavaraj A, Moraczewski D, Etzel JA (2023). Front. Neurosci. 17:1205928. [5] Fonov V, Evans AC, Botteron K, Almli CR, McKinstry RC, Collins DL, et al. (2011).

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Conclusions

While both alignment programs typically produce warped anatomicals that closely match the template structure, the new *sswarper2* provides more robust alignment across a broader range of datasets. AFNI's *sswarper2* results can also be integrated directly into *afni_proc.py*⁶ and other FMRI pipeline tools.

Acknowledgments, References & QR Code

To compare overall robustness of the programs, we tested each on a set⁴ of 169 anatomical T1w datasets from 8 different sites from 3 continents, with a wide subject age range (8-70 yrs), aligning to the MNI 2009c asymmetric template⁵. We compare individual and group-wide results both qualitatively and quantitatively.

The primary nonlinear alignment program in AFNI is *3dQwarp*² . *@SSwarper* wraps around *3dQwarp* and also includes removal of nonbrain material ("skullstripping"), since warping and skullstripping are each complicated processes that improve when the other is done well. If the skull+nonbrain region of a subject's anatomical dataset has been exactly removed, alignment to a template is much easier; or if a dataset has been well aligned to a template, one can use the latter to "punch away" the skull of the former. *@SSwarper* makes use of this by iterating between these steps with increasing accuracy, improving each. During alignment, it uses a local Pearson cost to drive early stages of alignment³. It also saves snapshots of initial and final stages of overlap, to provide quality control (QC) checks.

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Figure 1. Examples of anatomical-to-template alignment for (old) *@SSwarper* and (new) *sswarper2*. The underlay (grayscale) is the warped anatomical volume, and the overlay shows the edges of the MNI template. Pink arrows highlight local misalignments from *@SSwarper* that are correctly provided by *sswarper2*. While *@SSwarper* typically produces good alignment, outlier misalignments such as shown occur in a small fraction of cases. The *sswarper2* program is much more robust against these kinds of errors.

A) Mean of all @SSwarper and sswarper2 alignments to MNI

B) Difference in std dev of all @SSwarper and sswarper2 alignments to MNI

Figure 2. Summary group results of alignment. A) The crossgroup mean is shown for *@SSwarper* (top) and *sswarper2* (bottom); the edges of the reference MNI template are overlaid. In each case, the mean shows a fairly uniformly accurate alignment. B) The difference standard deviation (*@SSwarper* – *sswarper2*) of all warped anatomical datasets. Outside the brain, *@SSwarper* has higher values, since some alignments overstretched, while *sswarper2* tends to avoid such issues.

Similarly, *sswarper2* iterates between (local Pearson) alignment to a template and skullstripping, but it does so in smaller steps, more finely interleaved with skull removal, increasing stability. Additionally, *sswarper2* saves a detailed history of snapshots of intermediate processing steps, to facilitate any troubleshooting.

In many cases, *@SSwarper* and *sswarper2* yield quite similar results, providing accurate skullstripping and nonlinear registration to the MNI template. However, in a small number of cases, *@SSwarper* had inaccurate final skullstripping, resulting in localized misalignment. Fig 1 shows two examples of datasets in which the warped subject anatomy extends 2-3 mm outside the brain locally. In each case, *sswarper2* provides more accurate alignment and skullstripping. Fig 2A shows the mean of all warped datasets (each had been unifized to similar brightness values per tissue class) for each program. There is overall good alignment to the template in both cases. Fig 2B shows the voxelwise standard deviation across the group for each program. Again, overall patterns are similar, but *sswarper2* results are uniformly more tightly aligned within the brain volume and at major tissue boundaries, while the effect of the small fraction of stretched *@SSwarper* results can be seen around the brain edge.

A) sub-512: @SSwarper and sswarper2 alignments to MNI

B) sub-120: @SSwarper and sswarper2 alignments to MNI

Nonlinear alignment/registration is a common processing step in FMRI analysis. However, no alignment algorithm is perfect, and developers continue looking for ways to improve algorithms (subject to mathematical and practical constraints).