Automatic segmentation of the striatum and globus pallidus using MIST: Multimodal Image Segmentation Tool

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1 Segmentation examples in individual subjects

Example segmentation results for the left putamen in five subjects from the HCP80 dataset are shown in Fig. S1. The segmentation procedure used the $T_1$-weighted, $T_2$-weighted and FA images, but only the $T_1$-weighted volume is used for visualisation. Results for the combination of the caudate nucleus and nucleus accumbens are shown in Fig. S2. In this case, $T_1$- and $T_2$-weighted images were used for segmentation, as these contained adequate contrast. In addition, as the ventricles are invisible on an FA image, the corpus callosum represented the closest image edge for parts of the caudate nucleus; this introduced an ambiguity in finding the actual boundary of the caudate nucleus.

Results for the globus pallidus in the same dataset are displayed in Fig. S3. This is an example of a structure where there is no adequate contrast in the $T_1$-weighted volume. In the coronal slices, the superior boundary is clearly visible in the $T_2$-weighted volume. The medial boundary is not as clear on this volume and appears to be detected mainly on the basis of the FA-contrast.

Figure S4 shows a qualitative comparison of MIST with the mesh-based output from FIRST. MIST compares favourably and clearly improves on the output from FIRST in a number of areas. The segmentation of the globus pallidus illustrates the advantage of multimodal segmentation; FIRST fails to estimate the medial boundaries of the structure, whereas the added contrast in the $T_2$-weighted and FA images allow the new method to correctly estimate all boundaries.

Examples of the structures that were segmented in the 7T dataset are shown in Figure S5. The additional contrast offered by the high resolution $T_2$-weighted images, as well as the QSM ones, offers a clear advantage over segmentation based on just the $T_1$-weighted volume.
Putamen in five subjects (HCP80)

<table>
<thead>
<tr>
<th>Axial</th>
<th>Coronal</th>
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<tbody>
<tr>
<td><img src="image1.png" alt="Axial Slices" /></td>
<td><img src="image2.png" alt="Coronal Slices" /></td>
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<td><img src="image3.png" alt="Axial Slices" /></td>
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<td><img src="image5.png" alt="Axial Slices" /></td>
<td><img src="image6.png" alt="Coronal Slices" /></td>
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</tbody>
</table>

Figure S1: Segmentation of the left putamen in the HCP80 dataset in five subjects. Top three rows: axial slices, bottom three rows: coronal slices.
Caudate and accumbens in five subjects (HCP80)

499566 654754 672756 751348 756055

Figure S2: Segmentation of the left caudate nucleus and nucleus accumbens in the HCP80 dataset in five subjects. Top two rows: axial slices, middle two rows: sagittal slices, bottom two rows: coronal slices.
Figure S3: Segmentation of the left globus pallidus in the HCP80 dataset in five subjects, overlayed on the $T_1$-weighted, $T_2$-weighted and FA volumes. Top three rows: axial slices, bottom three rows: coronal slices. The putamen is shown in blue for reference.
Comparison between MIST and FIRST

<table>
<thead>
<tr>
<th>Subject</th>
<th>Putamen</th>
<th>Globus pallidus</th>
</tr>
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<tbody>
<tr>
<td>499566</td>
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<td>Axial</td>
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<tr>
<td>499566</td>
<td>Coronal</td>
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<td>899855</td>
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</table>

Figure S4: Comparison between MIST (red) and FIRST (blue) of segmentations of the putamen and globus pallidus in ten subjects of the HCP80 dataset, displayed on the $T_2$-weighted volume. Columns 1 and 3: axial slices, columns 2 and 4: coronal slices.
Example segmentations in 7T dataset

<table>
<thead>
<tr>
<th></th>
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<tbody>
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<td>T2*</td>
<td>QSM</td>
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<td>Putamen</td>
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<tr>
<td>Caudate + accumbens</td>
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<tr>
<td>Globus pallidus</td>
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</tbody>
</table>

Figure S5: Segmentation of the putamen (left: axial, right: coronal), the caudate nucleus and nucleus accumbens (left: sagittal, right: coronal) and the globus pallidus (left: axial, right: coronal) in one subject in the 7T dataset, overlaid on the $T_1$-weighted, $T_2^*$-weighted and QSM volumes.
2 Details of the intensity model

Logarithm of the probability and its derivatives  The logarithm of the probability Eq. 17 itself is

\[
\log p(M, L, \theta, \delta_1, \ldots, \delta_S | z_1, \ldots, z_S) = \text{const} + \sum_{m=0}^{N_r} \sum_{r=0}^{N_r} \log N(\mu_{mr} | \mu_{mr}^0, n_0 \Lambda_{mr})
\]

\[
+ \log \text{Wi}(\Lambda_{mr} | \alpha^0, \beta^0_{mr}) + \sum_{m=0}^{N_m} \log \text{Dir}(\theta_m | \alpha)
\]

\[
+ \sum_{s=0}^{N} \log \left( \sum_{\delta=0}^{\Delta} N(\delta | \mu, \lambda) \prod_{m=0}^{N_m} \prod_{r=0}^{N_r} N(z_{sm} | \mu_{mr}, \Lambda_{mr}) \text{Cat}(r | \theta_m) \right). (1)
\]

The derivatives with respect to its parameters are

\[
\frac{d}{d\theta_{nj}} \log p(M, L, \theta, \delta_1, \ldots, \delta_S | z_1, \ldots, z_S) =
\]

\[
\sum_{s=0}^{N} Z_s \left[ \sum_{\delta=0}^{\Delta} N(\delta | \mu, \lambda) \left( \prod_{m=0}^{N_m} \prod_{r=0}^{N_r} N(z_{sm} | \mu_{mr}, \Lambda_{mr}) \text{Cat}(r | \theta_m) \right) \right]
\]

\[
\times \left( N(z_{sn} | \mu_{nj}, \Lambda_{nj}) - N(z_{sn} | \mu_{nk+1}, \Lambda_{nk+1}) \right)
\]

\[
+ \frac{1 - \alpha_{k+1}}{1 - \sum_{\ell=1}^{k} \theta_{n\ell}} + \frac{\alpha_j - 1}{\theta_{nj}} (2)
\]

for the mixing coefficients,

\[
\frac{d}{d\mu_{nj}} \log p(M, L, \theta, \delta_1, \ldots, \delta_S | z_1, \ldots, z_S) =
\]

\[
\sum_{s=0}^{N} Z_s \left[ \sum_{\delta=0}^{\Delta} N(\delta | \mu, \lambda) \left( \prod_{m=0}^{N_m} \prod_{r=0}^{N_r} N(z_{sm} | \mu_{mr}, \Lambda_{mr}) \text{Cat}(r | \theta_m) \right) \right]
\]

\[
\times \text{Cat}(j | \theta_n) N(z_{sn} | \mu_{nj}, \Lambda_{nj}) \text{U}_\delta \left( (z_{sn} - \mu_{nj})^T \Lambda_{nj} \right)
\]

\[
- n_0 (\mu_{nj}^T - \mu_{nj}^0 \Lambda_{nj}) \Lambda_{nj} (3)
\]
for the component means and

\[
\frac{d}{d\Lambda_{nj}} \log p(M, L, \delta_1, \ldots, \delta_S | z_1, \ldots, z_S) = \\
\sum_{s=0}^{N} \frac{1}{Z_s} \left[ \sum_{\delta=0}^{\Delta} \mathcal{N}(\delta, \mu_{\delta}, \lambda_{\delta}) \left( \prod_{n=0}^{N_m} \sum_{r=0}^{N_r} \mathcal{N}(z_{sn} | \mu_{\delta_{mr}}, \Lambda_{\delta_{mr}}) \text{Cat}(r | \theta_m) \right) \\
\times \text{Cat}(j | \theta_n) \mathcal{N}(z_{sn} | \mu_{\delta_{nj}}, \Lambda_{\delta_{nj}}) \frac{1}{2} U_{\delta} \left( \Lambda_{\delta_{nj}}^{-1} - (z_{sn} - \mu_{\delta_{nj}})(z_{sn} - \mu_{\delta_{nj}})^T \right) \right]
\]

\[
+ (\alpha - \frac{k}{2}) \Lambda_{\delta_{nj}}^{-1} - \beta^0 - \frac{n_0}{2} (\mu_{nj} - \mu_{n_0}) (\mu_{nj} - \mu_{n_0})^T
\]

for the component covariances. The derivatives with respect to the diagonal of the matrix \( D_{mr} \) as defined in the main text are given by the diagonal of

\[
G^{-1} \left( \frac{d}{d\Lambda_{nj}} \log p(M, L, \delta_1, \ldots, \delta_S | z_1, \ldots, z_S) \right) G^{-1}.
\]

In these derivatives,

\[
Z_s = \sum_{\delta=0}^{\Delta} \mathcal{N}(\delta, \mu_{\delta}, \lambda_{\delta}) \prod_{m=0}^{N_m} \sum_{r=0}^{N_r} \mathcal{N}(z_{sn} | \mu_{\delta_{mr}}, \Lambda_{\delta_{mr}}) \text{Cat}(r | \theta_m).
\]

The notation \( U_{\delta}(x) \) is used to indicate that the lower-dimensional vector \( x \) is zero-padded to match a longer vector in the way that is appropriate for displacement \( \delta \).
3 Details of the shape model

After the intensity models for all vertices have been trained, we need to train the shape part of the model. As explained in the main text, this is done using the displacements found in the training data. These are found by simply selecting the most probable integer displacements given the MAP estimates of the parameters of the intensity models. The shape model is a simple multivariate normal distribution and we can use the standard inferential process as given by, for example, Bernardo and Smith (1994) to train it. This implies that we need to use a Normal-Wishart prior for the mean and precision parameters of the distribution. We are interested in the posterior predictive distribution

\[ p(\delta|\delta_0, \ldots, \delta_N_t) = \int_{\mu^s} \int_{\Lambda^s} p(\delta|\mu^s, \Lambda^s)p(\mu^s, \Lambda^s|\delta_0, \ldots, \delta_N_t)d\mu^sd\Lambda^s \quad (6) \]

where \(\delta_0, \ldots, \delta_N_t\) denote the displacements observed in the training data and

\[ p(\mu^s, \Lambda^s|\delta_0, \ldots, \delta_N_t) = \frac{p(\delta_0, \ldots, \delta_N_t|\mu^s, \Lambda^s)p(\mu^s, \Lambda^s)}{p(\delta_0, \ldots, \delta_N_t)} \quad (7) \]

with the Normal-Wishart prior

\[ p(\mu^s, \Lambda^s) = N_{N_t}(\mu^s_0, n_0^s\Lambda^s)W_{N_t}(\Lambda^s|\alpha, \beta) \].

The standard result is:

\[ p(\delta|\delta_0, \ldots, \delta_N_t) = St_{N_t}(\delta|\mu^s_t, \Lambda^s_t, 2\alpha_t) \quad (8) \]

where \(N_t\) is the number of training datasets and with

\[ \mu^s_t = \frac{n_0\mu^s_0 + N_t\delta^T}{n_0 + N_t} \]

\[ \Lambda^s_t = \frac{n_0 + N_t}{n_0 + N_t + 1} \alpha_t(\delta^T)^{-1} \]

\[ \beta_t = \beta + \frac{1}{2} \left( S + \frac{N_t n_0}{N_t + n_0}(\mu_0 - \delta^T)(\mu_0 - \delta) \right) \]

\[ \alpha_t = \alpha + \frac{1}{2} N_t - \frac{1}{2}(N - 1) \]

where \(\delta^T = \frac{1}{N_t} \sum_{j=1}^{N_t} \delta^T_j\) and \(S = \sum_{j=1}^{N_t}(\delta^T_j - \delta^T)(\delta^T_j - \delta^T)^T\). As before, \(N\) is the number of vertices, i.e. the dimensionality of the distribution.

To produce a segmentation for a new dataset, we need to find the MAP estimate of \(\delta\). We use an approach similar to the iterated conditional modes algorithm (Besag, 1986). To find the MAP estimate for \(\delta\), we sequentially find maxima of the conditional distributions of its components \(\delta_i\) given the other components. This procedure is repeated until convergence. The conditional probability of the displacement at a single vertex is the product of the conditional distribution of the shape model given all other components and the intensity model for that vertex. Up to a multiplicative constant, the conditional probability for vertex \(i\) is

\[ p(\delta_i|\delta_{\text{ic}}, Y_i, M_i, L_i, \theta_i, \mu^s_i, \Lambda^s) \propto St(\delta_i|\mu_{ij|^i}, \lambda_{ij|^i}, \alpha_{ij|^i})p(Y_i|\delta_i, M_i, L_i, \theta_i) \quad (9) \]
where the subscript $i^c$ denotes all components except $i$ and the second factor on the right-hand side is the intensity model for vertex $i$. The parameters of this distribution are

$$
\mu_{i|i^c} = \mu_i - (\Lambda_{ii}^s)^{-1} \Lambda_{ii}^s (\delta_i - \mu_i^s) \tag{10}
$$

$$
\Lambda_{i|i^c}^s = \Lambda_{ii}^s \left[ \frac{\alpha + N - 1}{\alpha + (\delta_i - \mu_i^s)(\Sigma_{i|i^c}^s)^{-1}(\delta_i - \mu_i^s)^T} \right] \tag{11}
$$

$$
\alpha_{i|i^c} = \alpha + N - 1, \tag{12}
$$

where $\Sigma_{i|i^c}^s = ((\Lambda_s)^{-1})_{i|i^c}$. This distribution is a consequence of the conditional distributions of a subset of the multivariate Student distribution given the other part, as described in, for example, Bernardo and Smith (1994).

The intensity models are only defined for integer displacements. In practice, we evaluate both the shape and intensity model factors at the points corresponding to the integer values of $\delta_i$ and optimise discretely. There is a possibility that the final segmentation mesh self-intersects. As the reference mesh is constructed in such a way that there are no self-intersections, we remove them from the final result by incrementally deforming a mesh from the reference shape to the fitted mesh. At each small increment we use VTK’s `vtkSelectEnclosedPoints` filter to test, for each vertex, whether it can be updated without generating a self-intersection. If the proposed displacement moves the vertex inwards with respect to the local surface then we know that the new point should be on the inside of the undeformed mesh (the mesh as it was before applying this update). If this proposed location is reported as being outside the mesh then it must have passed through some other part of the mesh and generated a self-intersection, in which case this point is held fixed. Otherwise, the proposed update is applied. Self-intersections for vertices that are being moved outward with respect to the reference mesh are detected in a similar fashion, but rejected if the proposed location is inside the undeformed mesh.
References
