New York Brain Bank

Why label brains?

Labels visually augment anatomy

- teach brain anatomy
- guide neurosurgery

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Labels compartmentalize image data

- assign results to brain regions vs. voxels
- quantify data by brain region vs. voxels

Why label brains?

Labels visually augment anatomy

- teach brain anatomy
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Labels compartmentalize image data

- assign results to brain regions vs. voxels
- quantify data by brain region vs. voxels

Labels provide a common nomenclature

- compare individuals within and across studies
- communicate results with a common language

Occipital Lobe
Gyrus angularis
Precuneus/ Cingulate cortex
Ventral striatum
Orbitofrontal Cortex

How should we label brain images?

Manual labeling (aMRI)

Atlas registration-based labeling (atlas)

Functional Mapping (fMRI)

Tractography-based segmentation (dMRI)

Feature matching (Mindboggle)

aMRI   atlas   fMRI   dMRI   Mindboggle

aMRI   atlas   fMRI   dMRI   Mindboggle

aMRI   atlas   fMRI   dMRI   Mindboggle

Cytoarchitectonic divisions

Korbinian Brodmann
(1868-1918)

aMRI   atlas   fMRI   dMRI   Mindboggle

Abb. 3.

Cytoarchitectonic divisions

Constantin von Economo
(1876-1931)

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aMRI atlas fMRI dMRI Mindboggle

Sir Grafton Elliot Smith
(1871-1937)

Topographic divisions

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aMRI   atlas   fMRI   dMRI   Mindboggle

Sulcus definitions

aMRI atlas fMRI dMRI Mindboggle

SFG   SPL

MFG   PrG   PoG   SMG   AnG   SOG

OpIFG   occ-ant   MOG

TrIFG   STG

OFC   OrIFG

MTG   IOG

ITG

Gyrus definitions

aMRI   atlas   fMRI   dMRI   Mindboggle

Gyrus definitions

aMRI   atlas   fMRI   dMRI   Mindboggle

Volume vs. surface label definitions

aMRI   atlas   fMRI   dMRI   Mindboggle

Volume vs. surface label definitions

aMRI atlas fMRI dMRI Mindboggle

Manual labeling software

aMRI　　　atlas　　　fMRI　　　dMRI　　　Mindboggle

Manual editing of cortical surface labels

aMRI   atlas   fMRI   dMRI   Mindboggle

Manual editing of cortical surface labels

aMRI   atlas   fMRI   dMRI   Mindboggle

Manual editing of cortical surface labels

Manual editing of cortical surface labels

Online brain image viewer

Online brain image viewer

An interactive tool for constructing optimal brain colormaps

Arno Klein
NY State Psych. Inst.
Columbia University, NY

Andrew Worth
Neuromorphometrics, Inc., MA

Jason Tourville
Cognitive & Neural Systems
Boston University, MA

Bennett Landman
Vanderbilt University
Nashville, TN

Tito Dal Canton
NY State Psych. Inst.
Columbia University, NY

Satrajit S. Ghosh
Research Laboratory of
Electronics, MIT, MA

David Shattuck
Laboratory of Neuroimaging
UCLA, CA

Goal

Create a software tool to aid in the construction of optimal colormaps for viewing and labeling brain anatomical images.

Background

The application of color is an indispensible means of visually distinguishing anatomical regions of the brain [1,2]. The need to facilitate visualization of brain images has become particularly urgent with the inception of large-scale anatomical labeling projects, such as the BrainCOLOR project, in which hundreds of MR images of the human brain are being manually labeled (www.braincolor.org/protocols) as well as viewed online (www.braincolor.org/roygbiv).

Choosing a color space

Our colormap must consist of colors that are perceptually distinct, so it is reasonable to choose a color space that is perceptually uniform, where a separation between two colors in the color space is as discriminable as the same separation between two other colors in the color space.
The International Commission on Illumination (CIE) devised a color space in 1931 that facilitates color description (Fig.1). Subsequent improvements led to the CIELAB and CIELUV color spaces, as well as a cylindrical variant, CIELch, all of which attempt to be perceptually uniform.

Fig.1

Constructing a colormap

We uniformly sampled hues from the CIELch cylinder (Fig.2) at different distances from the center (70-100% chromanance) and at different lightness values. The resulting colormap (Fig.3) has the same number of colors as regions we wish to color (49 cortical regions in the BrainCOLOR protocol). We intend to expand the number of colors to include subcortical and other regions.

Fig.2

Fig.3

1. Schott GD (2010) Colored illustrations of the brain: some conceptual and contextual issues. Neuroscientist. 16(5):508-18.
2. Schoonover, C (2010) Portraits of the Mind: Visualizing the Brain from Antiquity to the 21st Century. Abrams (ISBN-13: 978-0810990334)

CIE1931 figure: http://en.wikipedia.org/wiki/File:CIExy1931_MacAdam.png
CIELch figure: http://www.colorbasics.com/ColorSpace/

Maximizing discriminability of adjacent regions

To easily distinguish brain regions from each other, we wish to maximize the color difference between adjacent regions. We do this by constructing a graph that represents connections (edges) between regions (nodes).

We define an *optimal mapping* of colors as the combination of colors assigned to nodes with the maximum sum of color differences between connected nodes, defined by the CIE2000 Delta E measure (http://en.wikipedia.org/wiki/Color_difference).
However, we cannot compute 8.8×10^{18} permutations of the 49 colors in the colormap, so we break up the brain into sublobes and compute the permutations of portions of the colormap at a time:

Frontal Lobe (7,6,9):
- Lateral Surface (7)
- Medial Surface (6)
- Inferior Surface (4)
- Opercular Region (3)
- Insular Region (2)

Temporal Lobe (8):
Lateral Surface (4)
Supratemporal Surface (3)
Inferior Surface (1)

Parietal lobe (6):
Lateral Surface (4)
Medial Surface (2)

Occipital Lobe (8):
Lateral Surface (4)
Inferior Surface (1)
Medial Surface (3)

Limbic Cortex (5):
Cingulate Cortex (3)
Medial Temporal Cortex (2)

Grouping regions by using similar colors

For labeling a brain image, maximum discriminability is paramount. For viewing a brain image, however, it is useful to visually group regions by assigning similar colors to the regions within a group. To create this visual grouping, we colored each subgraph with a neighborhood of colors (sampled within a wedge of the CIELch cylinder).

7 color groups; lightness: 60, 75, 90; chromaticity: 70%

7 color groups; lightness: 60, 75, 90; chromaticity: 100%

15 color groups (sublobes); lightness: 50, 70, 90; chromaticity: 70%

Conclusion and Future Work

We have created Python software for creating and applying colormaps to brain images.
Download the software from: http://binarybottle.github.com/brainCOLORmap
The software calls the Python libraries: *NumPy, NetworkX, Python-Colormath, Matplotlib, and xlrd,* to:

1. convert a table of adjacent regions (an adjacency matrix) to a weighted graph of colored nodes
2. compute permutations of colors for each subgraph to optimally distinguish colors of adjacent regions
3. apply similar colors to predefined groups of regions

There are many improvements we could pursue. We have targeted the following for future work:

➜ Develop an optimization algorithm for whole graphs or larger subgraphs.
➜ Restrict the color space to colors that can be displayed; currently, the CIELch colors that should appear different from one another may appear similar after conversion to SRGB colors.
➜ Seed color values to better control the range of colors for each group of regions.
➜ Accommodate visual concerns such as color blindness by adapting the colormap accordingly.

Optimal brain colormaps

aMRI **atlas** fMRI dMRI Mindboggle

atlas   target

Atlas registration-based labeling

aMRI   atlas   fMRI   dMRI   Mindboggle

atlas   target

transform

Atlas registration-based labeling:

Step 1:   compute the registration transform
              from the atlas to the target

aMRI   atlas   fMRI   dMRI   Mindboggle

atlas   target

transform

Atlas registration-based labeling:

Step 2:   apply the transform to the atlas labels

aMRI   atlas   fMRI   dMRI   Mindboggle

atlas   target

transform

Atlas registration-based labeling:

Step N: compute and apply transforms for multiple atlases

Volume- vs. surface-based registration

aMRI   atlas   **fMRI**   dMRI   Mindboggle

R   L

-8   ±2.3   +8

aMRI · atlas · **fMRI** · dMRI · Mindboggle

Visuotopic mapping

aMRI   atlas   fMRI   dMRI   Mindboggle

Functional "connectivity"
7-network (N=1000)

aMRI   atlas   fMRI   dMRI   Mindboggle

Purple (Visual)
Blue   (Somatomotor)
Green   (Dorsal Attention)
Violet (Ventral Attention)
Cream   (Limbic)
Orange (Frontoparietal)
Red   (Default)

Functional "connectivity"
7-network (N=1000)
Yeo, et al., JNeurophysiology, 2011

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aMRI   atlas   fMRI   dMRI   Mindboggle

Purple (Visual)
Blue   (Somatomotor)
Green   (Dorsal Attention)
Violet (Ventral Attention)
Cream   (Limbic)
Orange (Frontoparietal)
Red   (Default)

Functional "connectivity"
17-network (N=1000)
Yeo, et al., JNeurophysiology, 2011

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aMRI   atlas   fMRI   dMRI   Mindboggle

L   R   L   R

y=10   x=50   z=35

Functional "connectivity"

aMRI   atlas   fMRI   **dMRI**   Mindboggle

Visualization of atlas-based labels + tractography

aMRI   atlas   fMRI   dMRI   Mindboggle

LH   RH

LH   RH

Labeled tractography networks

Hagmann, et al., 2008

aMRI   atlas   fMRI   dMRI   Mindboggle

Tractography + atlas-based labeling

Wiegall et al., 2003; Cachia et al., 2008

aMRI   atlas   fMRI   dMRI   Mindboggle

Unimodal Mindboggle

aMRI atlas fMRI dMRI Mindboggle

aMRI

aMRI   atlas   fMRI   dMRI   Mindboggle

Depth map

aMRI   atlas   fMRI   dMRI   Mindboggle

Medial surfaces of folds

aMRI   atlas   fMRI   dMRI   Mindboggle

Folds (sulci)

48

aMRI   atlas   fMRI   dMRI   Mindboggle

Folds on an inflated surface

aMRI atlas fMRI dMRI Mindboggle

basins

ribbons

curves

points

Nested features

aMRI   atlas   fMRI   dMRI   Mindboggle

basins
volume
surface area
cortical thickness / sulcal span
...

ribbons
convex hull volume
surface area
depth
curvature
convexity
...

curves
length / tortuosity
depth
curvature
convexity
...

points
convex hull volume
sequence along a curve
...

Shape analysis

aMRI   atlas   fMRI   dMRI   Mindboggle

2nd spectra   6th spectra   8th spectra

Global to local

LBO on sulcal basin   LBO on sulcal basins of left hemisphere

Shape analysis: Laplace-Beltrami spectra

aMRI   atlas   fMRI   dMRI   Mindboggle

Ribbon (R)

Basin (B)

Fundus (F)

Pit (P)

Features within a subfold

aMRI   atlas   fMRI   dMRI   Mindboggle

Surface patches

Sulcus folds + subfolds

Structures within a subfold

Ribbon (R)

Basin (B)

Fundus (F)

Pit (P)

Feature graph model

B ← R ← F

Has label L

Part of

P1 ⟷ P2 ⟷ P3 ⟷ P4

Connected to

Nested features within a graph-based data model

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Graph-based database of hierarchical brain features
finding similar features

aMRI atlas fMRI dMRI Mindboggle

basin 88

basin 89

CONTAINS CONTAINS CONTAINS CONTAINS CONTAINS

86: basin 87: basin 91: basin 90: basin

88: basin IS SIMILAR IS SIMILAR 89: basin

HAS LABEL

75: precuneus 85: insula 78: superiorfrontal 67: paracentral 73: posteriorcingulate 64: medialorbitofrontal 76: rostralanteriorcingulate 82: frontalpole

62: lateralorbitofrontal 52: caudalanteriorcingulate 60: isthmuscingulate

Graph-based database: example query

aMRI   atlas   fMRI   dMRI   Mindboggle

Data layer

Mindboggle analytics engine

Web interface

T1-MRI

FreeSurfer (FS)
Surfaces | Labels

noSQL Graph Model

reXster
Gremlin
Pipes Frames
Blueprints

Graph Database
Neo4J

HTTP /
REST

NestedStructureExtractor
Pits | Fundi | Basins

ShapeCharacterization
Geometric | Shape | Spectral

DataConverter
JSON | Three.js | VTK | FS | GraphML

GraphDataImporter
Shapes | Features | Labels | Relations

Backend ↔ MVC ↔ Frontend

Server
NodeJS
Apache Lucene
Tinkerpop

Java
Groovy
Python
Jython

Visualization
WebGL
Three.js

User Interface
ExtJS
jQuery
HTML5
CSS3
JavaScript

The Mindboggle graph-based database infrastructure showing the individual components of the data layer, the analytics engine, and the web interface.

The graph model can be queried via REST, Cypher, reXster, or Gremlin.
This example shows how to query all outgoing relations from basin 88:

Gremlin query:
```
{
"script": "g.v(88).outE"
}
```

JSON response:

```
[{
  "start" : "http://192.168.23.219:7474/db/data/node/88",
  "data" : {
  },
  "type" : "IS SIMILAR",
  "extensions" : {
  },
  "end" : "http://192.168.23.219:7474/db/data/node/89"
}, {
  "start" : "http://192.168.23.219:7474/db/data/node/88",
  "data" : {
  },
  "type" : "HAS LABEL",
  "extensions" : {
  },
  "end" : "http://192.168.23.219:7474/db/data/node/60"
},
```

Graph-based database schematic

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aMRI atlas fMRI dMRI Mindboggle

whole brain

single folds

Basins Both Hemispheres Renderer

Folds Renderer

inflated pial white matter

aMRI   atlas   fMRI   dMRI   Mindboggle

atlas   target

aMRI   atlas   fMRI   dMRI   Mindboggle

Mindboggle step 1:   extract features, compute their shape measures

atlas   target

aMRI   atlas   fMRI   dMRI   Mindboggle

Mindboggle step 2:   match atlas and target features

atlas   target

match

aMRI   atlas   fMRI   dMRI   Mindboggle

Mindboggle step 3:   compute feature-driven transform from atlas to target

atlas   target

transform

aMRI   atlas   fMRI   dMRI   Mindboggle

Mindboggle step 4:   apply transform to labels

atlas   target

transform

aMRI   atlas   fMRI   dMRI   Mindboggle

OR   extract, match, and fill labels without registration!

atlas   target

extract   fill

match

aMRI   atlas   fMRI   dMRI   Mindboggle

Sulcal fundi

aMRI   atlas   fMRI   dMRI   Mindboggle

Sulcal fundi

aMRI atlas fMRI dMRI Mindboggle

Sulcal fundi + manual labels

aMRI   atlas   fMRI   dMRI   Mindboggle

Initial label propagation test: manual labels

aMRI   atlas   fMRI   dMRI   Mindboggle

Initial label propagation test: results

aMRI　　　atlas　　　fMRI　　　dMRI　　　Mindboggle

Initial label propagation test: rank label errors

aMRI   atlas   fMRI   dMRI   Mindboggle

atlas   target

extract   fill

match

aMRI   atlas   fMRI   dMRI   Mindboggle

Multimodal Mindboggle

aMRI   atlas   fMRI   dMRI   Mindboggle

L   R

y=10   x=50   z=35

fMRI "connectivity" clusters as features

aMRI   atlas   fMRI   dMRI   Mindboggle

Legend:
- CST
- FX
- CG
- SLF
- ARC
- IOF
- CC-FM
- UNC

dMRI tract clusters as features

aMRI   atlas   fMRI   dMRI   Mindboggle

dMRI tract clusters as features

aMRI   atlas   fMRI   dMRI   Mindboggle

dMRI tract clusters as features

aMRI   atlas   fMRI   dMRI   Mindboggle

fMRI-connected fMRI features: ADHD

aMRI   atlas   fMRI   dMRI   Mindboggle

fMRI-connected aMRI features: ADHD

aMRI   atlas   fMRI   dMRI   Mindboggle

dMRI-connected aMRI features: Major depressive disorder

aMRI   atlas   fMRI   dMRI   Mindboggle

A gray/white matter surface (left lateral view) with visible sulcal pits highlighted (cyan circles). These features go by different names (sulcal roots, buried or annectant gyrii, plis de passage) and may be well conserved structures formed early in development.

DTI connectivity graph (same subject). Vertices represent all extracted sulcal pits and each edge width indicates a connection probability greater than 0.01 between two vertices (and does not follow a tractography path).

dMRI-connected aMRI features: Major depressive disorder

aMRI · atlas · fMRI · dMRI · Mindboggle

Early attempt at subgraph extraction (upper row) and matching (lower row).
These graphs were constructed from (left to right) a remitter, non-remitter, and control subject.
The subgraphs in red in the lower row have the highest small-worldness ratio.

dMRI-connected aMRI features: Major depressive disorder

aMRI   atlas   fMRI   dMRI   Mindboggle

Clinical application: Prediction of treatment response based on feature shapes

We have encouraging preliminary evidence that we can predict treatment response for major depressive disorder based solely on fundus shape information! (Klein et al, SFN2011)

Greg Millington
Neuromorphometrics

Mindboggle team

aMRI surfaces

Joachim Giard
Univ. Catholique de
Louvain, Belgium

aMRI features

Forrest Bao
Texas Tech

fMRI features

Gael Varoquaux
NeuroSpin, France

dMRI features

Eleftherios Garyfallidis
Cambridge University

Feature matching

Yrjö Häme
Graduate student

**Shape analysis &
label propagation**

Eliezer Stavsky
Graduate student

Graph database

Nolan Nichols
Washington U.

Web visualization

Noah Lee
former postdoc

Example of natural morphological variability: left inferior parietal lobule (IPL; Kiriyama et al. 2009).
(A-D) are MRI data and (E-G) are post-mortem specimens.
(A) IPL is highlighted and folds are outlined.
(B,E) Typical folding pattern.
(C,F) PreSMG pattern: an additional gyrus (ellipse) lies between postCS and SMG.
(D,G) PreAG pattern: an additional gyrus (ellipse) lies between SMG and AG.

[SMG: supramarginal gyrus; AG: angular gyrus; postCS: postcentral sulcus; IPS: intraparietal sulcus;
Sy: Sylvian fissure, STS: superior temporal sulcus; *sulcus intermedius primus]

Characterize anatomical variability

Convert one brain graph into another!

Identify nested structures in our data