

## Automatic Segmentation of Traumatic Brain Injury MRI volumes using Atlas Based Classification and 3D Slicer

Andrei Irimia, Micah C Chambers, John D. Van Horn Laboratory of Neuro Imaging University of California, Los Angeles andrei.irimia@loni.ucla.edu 310-206-2101 http://loni.ucla.edu

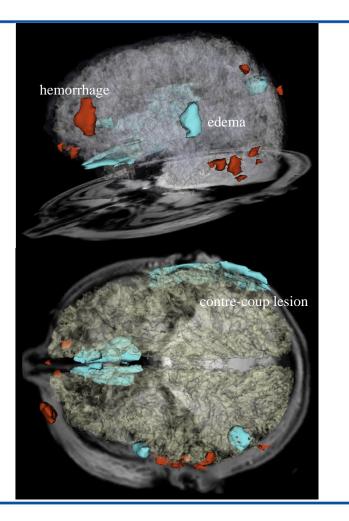
NA-MIC Tutorial Contest: Summer 2011



## This tutorial is being submitted to the End-to-End Solution contest category.

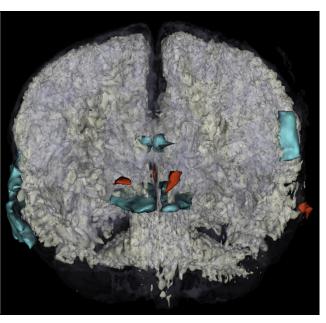


- acquire familiarity with several MR sequences commonly used for TBI imaging
- learn how to apply ABC to perform joint co-registration and automatic segmentation of TBI volumes
- acquire expertise on how to identify and characterize TBI pathology using various MRI sequences
- gain exposure to informed strategies for quantification of TBI-related edema or hemorrhage





- This tutorial assumes that you have already completed the following tutorials:
- Slicer 3 Visualization Tutorial by Sonia Pujol, available at
  - <u>http://www.slicer.org/slicerWiki/images/2/2e/Slicer3\_Data</u>
     <u>LoadingAndVisualization\_UCSF2010\_SoniaPujol.pdf</u>
- Interactive Editor Tutorial by Sonia Pujol, available at
  - <u>http://www.slicer.org/slicerWiki/images/6/69/InteractiveEditorT</u> <u>utorial\_Slicer3.6-SoniaPujol.pdf</u>



Edema (cyan) and hemorrhage (red) in a traumatic brain injury patient, with white matter (solid beige) and gray matter (transparent hue) superposed.

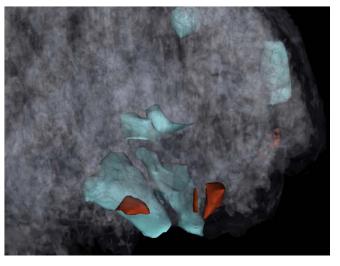


This tutorial requires the installation of the **Slicer3.6 release** and the tutorial dataset. They are available at the following locations:

Slicer3.6 download page

http://www.slicer.org/pages/Downloads/

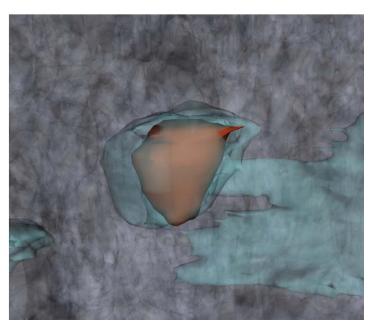
**Tutorial dataset:** The dataset can be downloaded <u>here</u>.



Enhanced view of frontal edema and hemorrhage several days after the insult. This type of pathology is very common in traumatic brain injury.



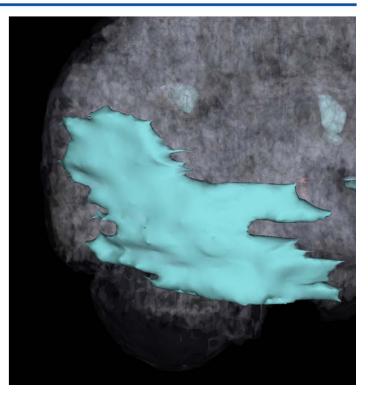
- This tutorial has been developed on Windows 7. It is compatible with the following platforms:
  - Windows XP, Windows 7
  - Linux 32, Linux 64
  - Mac/Darwin



Detailed view of parietal edema (teal) and hemorrhage. It is interesting to note how the non-hemorrhagic edemic region encases the brain portion that is actively bleeding. With recovery, it is expectable for the hemorrhage to shrink in both volume and mass.



- Clinical background
- Clinical workflow
  - installation of the ABC module
  - loading and exploration of TBI data
  - understanding common MR sequences for TBI
  - automatic segmentation of TBI using ABC
  - generation of 3D models
  - lesion segmentation and model
     generation
- Conclusions

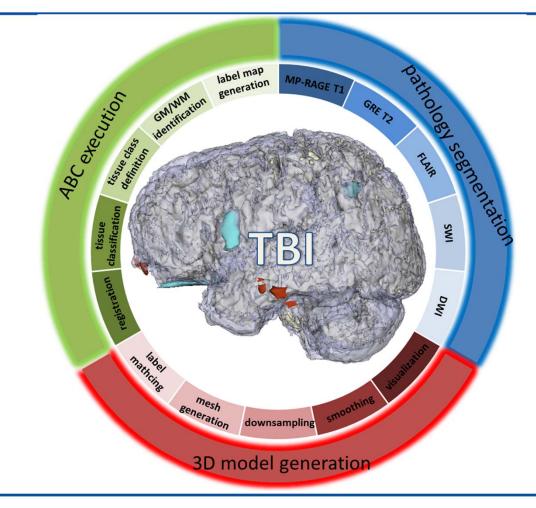


Contre-coup edema due to acceleration/deceleration forces at work during traumatic brain injury. Although contralateral with respect to the primary injury, the spatial extent of this edemic region is notable.

# Traumatic Brain Injury (TBI)

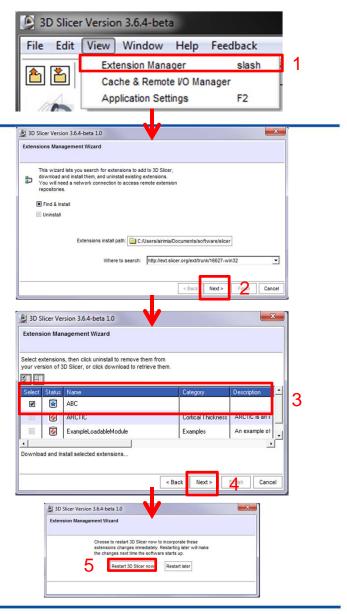
- With traumatic brain injury (TBI), there are an estimated 1.7 million cases in the United States alone every year, with an estimated 1.2 million ER visits and over 50,000 deaths.
- Over 5.3 million cases of required long-term daily assistance exist as a result of TBI, which results in health care costs of over \$60 billion/year
- Despite many innovations, progress towards patient-tailored characterization of the structural and functional substrates associated with TBI-related neural and cognitive impairment remains dissatisfactory and the relationship between neurophysiological markers of cognitive dysfunction and TBI structural damage has not been acceptably elucidated
- 3D Slicer offers a powerful and unparalleled set of tools for the exploration and quantification of TBI





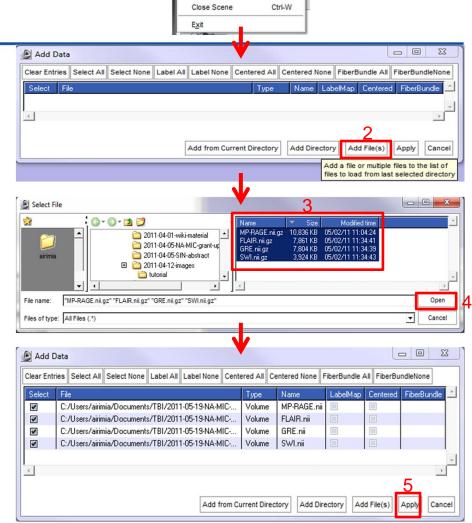


- After Slicer has been loaded, the Welcome window will appear.
- To install the ABC module:
  - 1 Click on View → Extension Manager
  - 2 Click on Next
  - 3 Select ABC from the list of modules
  - 4 Click on Next; you will be prompted to specify the local hard drive location for ABC
  - 5 After installation, you must select *Restart 3D Slicer now* for the changes to take effect





- After Slicer has been loaded, the Welcome window will appear.
- To load the TBI volumes associated with the case study:
  - 1 Click on *File* → *Add Data*
  - 2 Click on Add File(s)
  - 3 Navigate to the data folder and sort the volume files in descending order by size. This will ensure that all files are later processed in proper order.
  - 4 Click on Open
  - 5 In the Add Data dialog box, click on Apply



3D Slicer Version 3.6.4-beta
 File Edit View Window Help

CtrLO

Ctrl-A

Ctrl-S

Load Scene...

Import Scene.

Add Volume...

Save

Download Sample Da Add Data...

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- 4 TBI data volumes are provided for this tutorial: MP-RAGE, FLAIR, GRE, and SWI.
- MP-RAGE: this is an anatomical volume acquired using a T1-weighted sequence, in which white matter has higher intensity than gray matter
  - To explore a TBI-related lesion in this patient:
    - 1 link the views using S

    - 3 navigate to slice 112 using the slider (
  - The lesion associated with open-head TBI becomes apparent as a hypo-intensity located in the parieto-occipital region (see circle to the right)



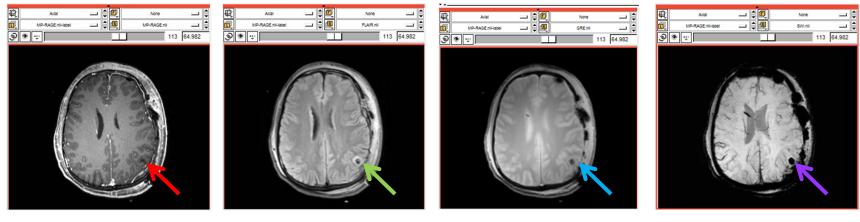


- To locate the lesion more accurately within the head:
  - 1 navigate to slice 179 in the sagittal view
  - 2 navigate to slice 79 in the coronal view
- The lesion and open head injuries are now apparent on all three views
- Similar exploratory navigation can reveal the lesion in FLAIR, GRE, and SWI volumes



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MP-RAGE T1

FLAIR

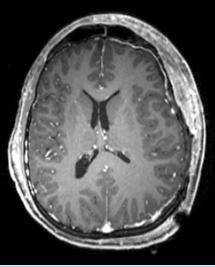
GRE T2

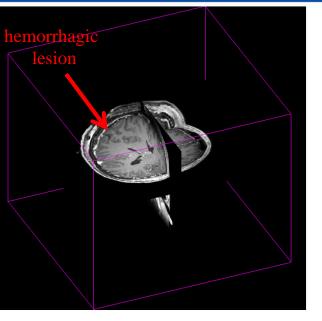
SWI

- Use of multiple image channels provides complementary information on pathologies
- T1 is a standard volume, with good anatomical resolution and high WM/GM contrast
- FLAIR is excellent for revealing CSF-perfused lesions as image hyper-intensities
- T2 GRE is useful for discerning areas associated with hemorrhages
- SWI is suitable for the detection of micro-bleeds and is superior to GRE in this respect



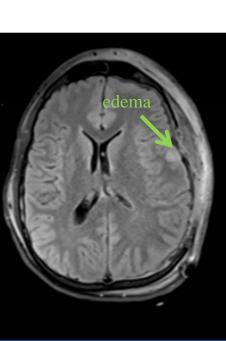
- MP-RAGE = magnetization prepared rapid gradient echo
- This sequence is a fast 3D gradient echo sequence designed for rapid acquisition with T1-weighted dominance, rapid sampling time, high signal intensity and contrast
- Although ideal for the acquisition of anatomical data sets, the abilities of this sequence to detect TBI-related pathology is very limited and its use must usually be complemented by other sequences

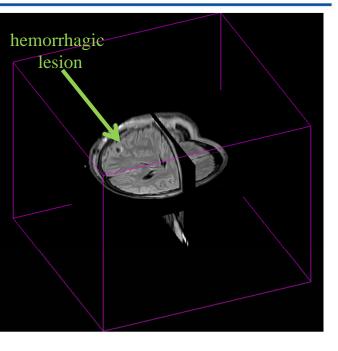


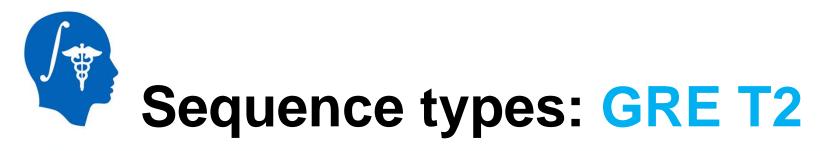




- FLAIR = fluid attenuation inversion recovery
- This sequence is an inversion recovery sequence with long TI adjusted to the TR of a fluid type whose effect is sought to be removed from the result image
- Hyperintensities in FLAIR images are indicative of CSF-perfused, edemic cortical tissue
- Compared to other sequences, FLAIR is not very useful for hemorrhage detection



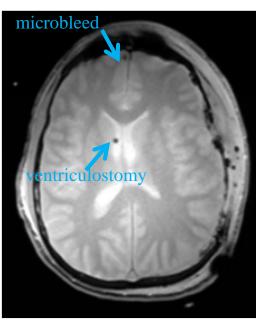


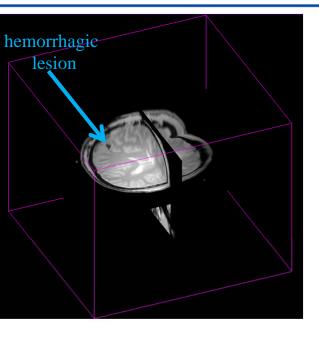


- GRE = gradient recalled echo
- This T2 weighted sequence has some very useful, though rather limited, ability to identify hemorrhage
- In our dataset, it can identify one microbleed in the

frontal lobe, as well as the ventriculostomy

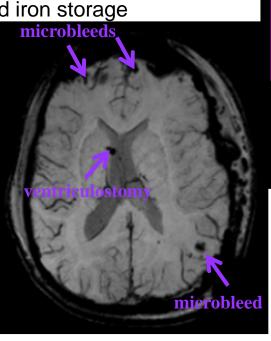
 Some studies have proposed that, although useful, GRE T2 sequences are not as sensitive to microbleeds as SWI

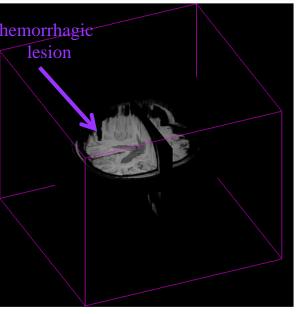






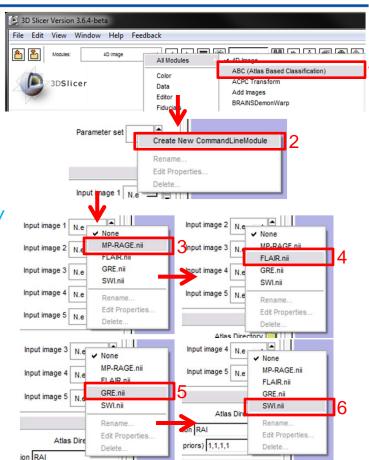
- SWI = susceptibility weighted imaging
- This sequence exploits susceptibility differences between tissues to create an enhanced contrast magnitude image which is exquisitely sensitive to venous blood, bleeds, and iron storage
- In our dataset, this sequence identifies several micro-bleeds as well as the ventriculostomy
- SWI is a classical sequence used for TBI imaging which can reveal the path of the ventriculostomy





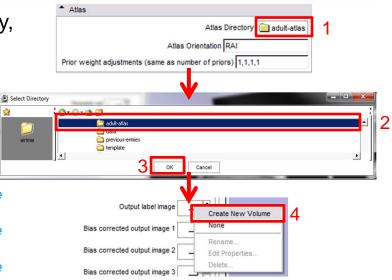


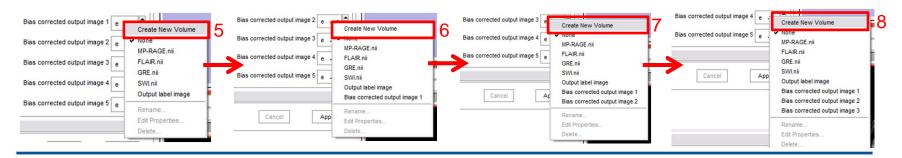
- To load ABC, click on:
  - 1 –Modules → All Modules → ABC
- The ABC interface appears. To specify input data:
  - 2 click Parameter Set → Create New Command Line Module
  - 3 click Input image 1 → MP-RAGE.nii
  - 4 click Input image 2 → FLAIR.nii
  - 5 click Input image 3 → GRE.nii
  - 6 click Input image 4 → SWI.nii





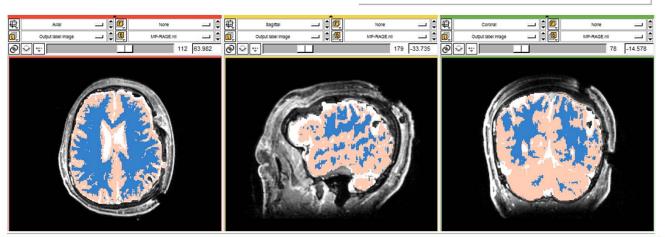
- 1 click on and locate the *adult-atlas* directory, which was included in the tutorial data set
- 2 select adult-atlas from its location
- 3 click OK to select the atlas template
- 4 click Output label image → Create New Volume
- 5 Click Bias corrected output image 1 → Create New Volume
- 6 Click Bias corrected output image 2 → Create New Volume
- 7 Click Bias corrected output image 3 → Create New Volume
- 8 Click Bias corrected output image 4 → Create New Volume

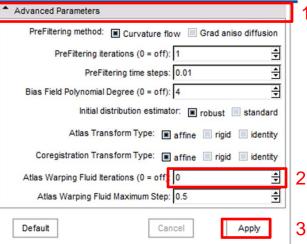






- 1 click on the Advanced Parameters arrow
- 2 enter 0 (off) for the number of fluid iterations
- 3 click *Apply* to run ABC on the dataset
- ABC will now co-register the FLAIR, GRE and SWI images to the MP-RAGE image, and perform the tissue classification
- Allow up to several hours for ABC execution
- ABC produces 3 tissue categories:
  - gray matter
  - white matter
  - CSF







Now that we have obtained the label map of GM, WM and CSF using ABC, we can generate the 3D models associated with each of these three tissue types.

- 1 From the Modules drop-down menu, select All Modules
- 2 From the second column, select Model Maker

This will display the interface of the *Model Maker* module, which allows one to create 3D models in Slicer.

3D Slicer Version 3.6.4-bet	ta		
File Edit View Window	Help Feedback		
Modules: ABC (A	All Modules	4D Image	Fast Rigid registration
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	Transforms	Cast Image	Gravscale Grind Peak
<ul> <li>Input Images</li> </ul>	VolumeRendering	ChangeTracker	Grayscale Model Maker
	Volumes	CheckerBoard Filter	Histogram Matching
	Wizards •	ClipModel	IA_FEMesh
	Informatics	CollectFiducials	IGTRecorder
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	Filtering •	CropVolume	Labelmap Seeding
	Surface Models	Curvature Anisotropic Diffusion	Linear registration
▲ Atlas	Converters +	Data	MRI Bias Field Correction
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B	as corrected output image	Extract Skeleton	NeuroNay
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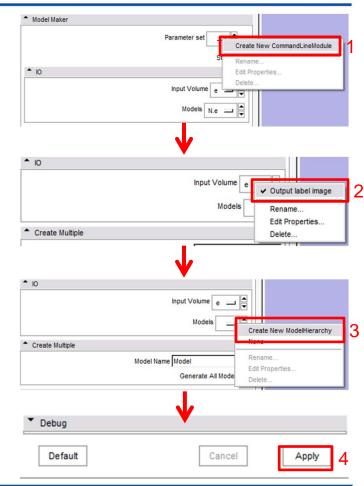


To generate the 3D models:

- 1 From the Parameter Set drop-down menu, select Create New Command Line Module
- 2 From the Input Volume drop-down menu, select Output label image
- 3 From the *Models* drop-down menu, select *Create New Model Hierarchy*

• 4 – Click Apply

These steps will create 3D models for WM, GM and CSF.

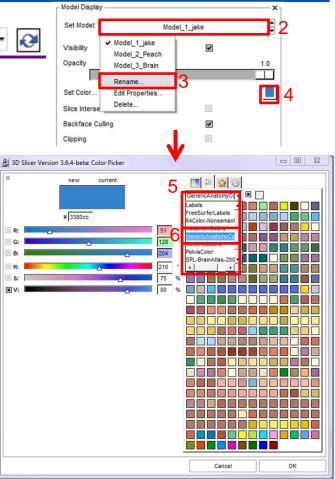






- 1 From the tool bar, click on the *Models* button
- 2 From the Set Model drop-down menu, select Model\_1\_jake
- 3 From the Set Model drop-down menu, select Rename and type WM; repeat steps 1-2 to rename Model\_2\_Peach as GM and Model\_3\_Brain as CSF
- 4 Set the model to WM as in step 2, and click on the color box;
- 5 In the Color Picker dialog box, click on the color drop down menu
- 5 Select Generic Anatomy Colors as the color map; this will assign a color scheme in accordance with Slicer conventions.





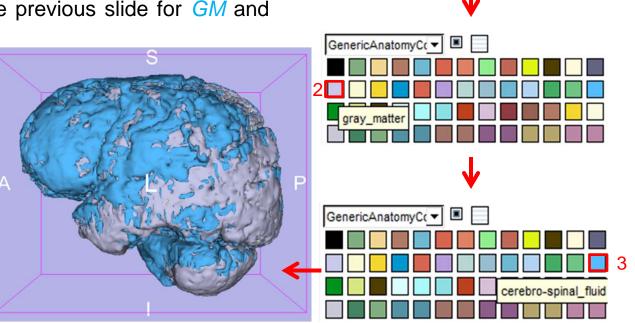


Now that we have selected an appropriate color scheme, we need to assign suitable colors to each tissue type.

 1 – In the Color Picker dialog box, select beige for WM

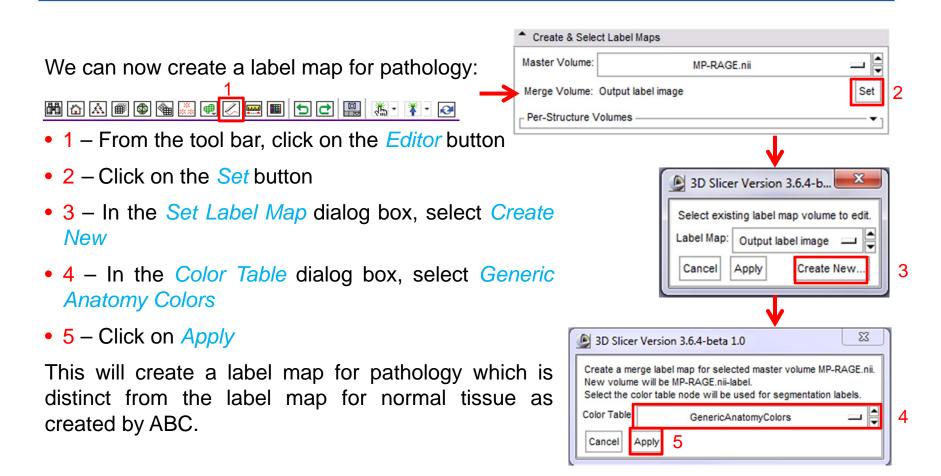
Repeat steps 2-5 on the previous slide for *GM* and *CSF* 

- 2 For *GM*, click on purple as shown
- 3 For *CSF*, click on blue as shown. The 3D model should now look as to the right. Note the variable CSF thickness.



GenericAnatomyC( -

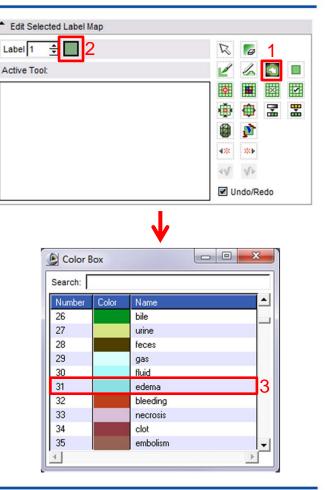






Now we can assign labels for the two pathology types, namely edema and bleeding:

- 1 From the label edit bar, click on *Level Tracing*
- 2 Click on the color button next to the Label
- 3 In the Color Box dialog box, select 31: edema





- 1 In the Foreground menu, select FLAIR.nii
- 2 Click on the tool bar menu
- 3 select Red slice only

#### Layout

 4 – Use the level tracing tool to segment the edema using the FLAIR volume.

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Toggle GUI panel visibility

Toggle GUI panel L/R

close

Recall that edema is hyperintense in this sequence modality, and that it appears in slices 105-117. The hypointense region in the middle is actually hemorrhage, but it will be segmented separately in following steps so we need not worry about it for now.

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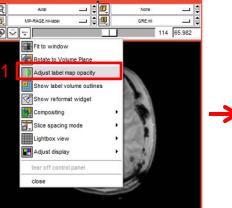


Once the edema has been segmented, we can proceed to the segmentation of hemorrhage. Viewing the label map and the underlying image at the same time can be accomplished by adjusting label map opacity.



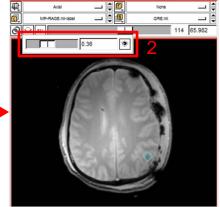
- 2 Use the slider to select a convenient opacity
- 3 In the Color Box dialog box, select 32: bleeding

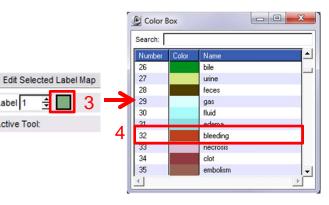
This step ensures that 3D Slicer color labeling conventions are respected.



Label 1

Active Tool:





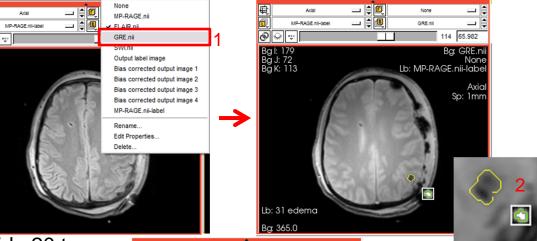


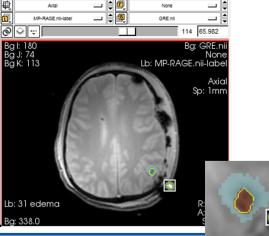
We can now segment bleeds.

- 1 From the red slice background menu, click on GRE.nii
- 2 Use the label map opacity slider as shown on the previous slide as well as the level

tracing tool as demonstrated on slide 20 to obtain a convenient view of the hemorrhage in relation to the surrounding edema.

3 – Segment the bleeding in slices 106-116. The label map should now be similar to the inset shown to the right. Note that hemorrhage is surrounded by edemic region which does not bleed.





# Create pathology 3D models

Once pathology has been segmented, their 3D models can be created.

• 1 - From the tool bar menu, select

the Conventional layout

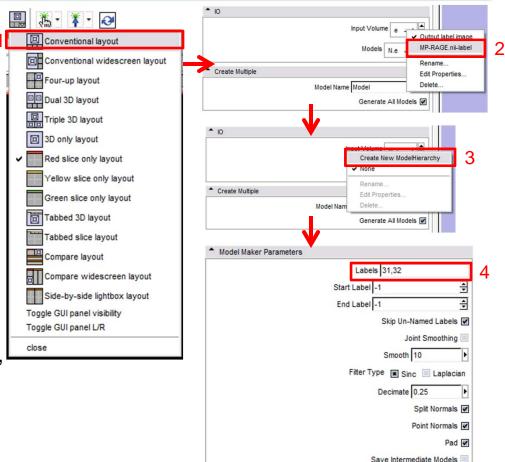
• 2 - For Input Volume, select MP-

*RAGE.nii-label.* This is the label map for the pathology.

• 3 – Under Models, select Create

### New Model Hierarchy

- 4 Under Model Maker Parameters,
- type "31,32" in the *Label* input box.
- 5 Click on Apply





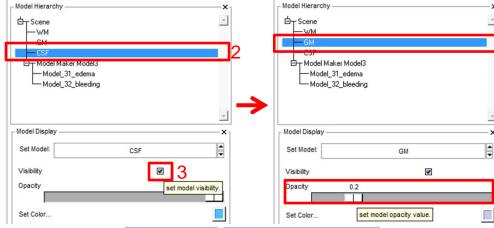
Hierarchy & Display

We can now visualize the full model, which includes both pathology and

healthy-looking tissue.

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- 1 On the tool bar, click Models
- 2 Under *Hierarchy and Display,* select *CSF*

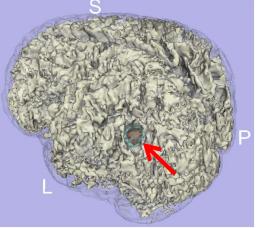


Hierarchy & Displa

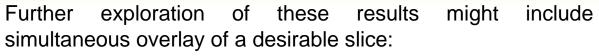
5

- 3 Under *Visibility*, uncheck the toggle box
- 4 Under Hierarchy and Display, select GM
- 5 Decrease the opacity to 0.2

This will allow us to visualize the edema and hemorrhage in relationship to the rest of the brain, as shown to the right.





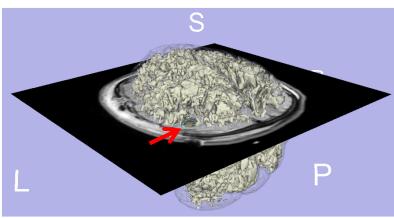


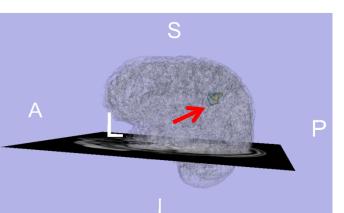
• 1 – On the red box tool bar, toggle slice visibility (left below)

One can also display the pathology with GM/WM transparency (see figure to the right below)

• 2 – Under Hierarchy and Display, set WM opacity to 0.2





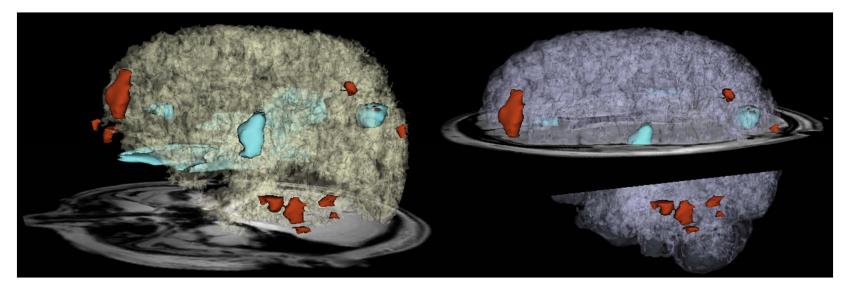


National Alliance for Medical Image Computing http://www.na-mic.org

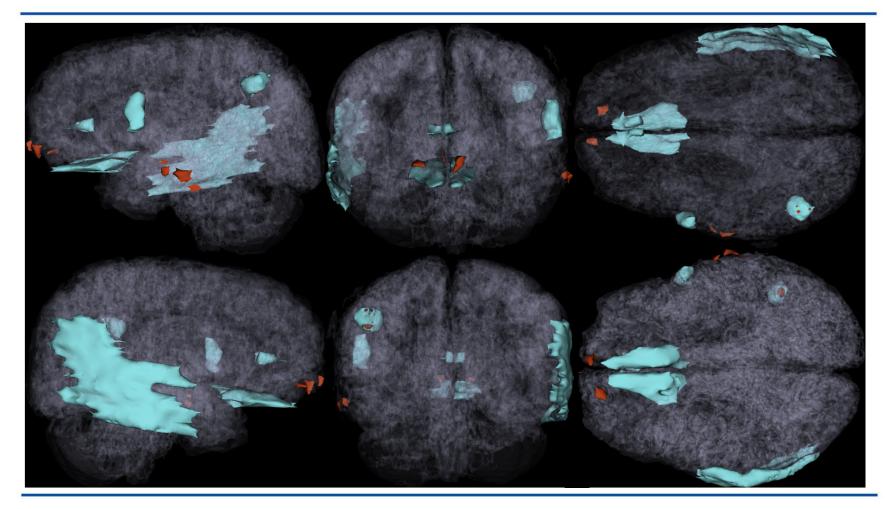
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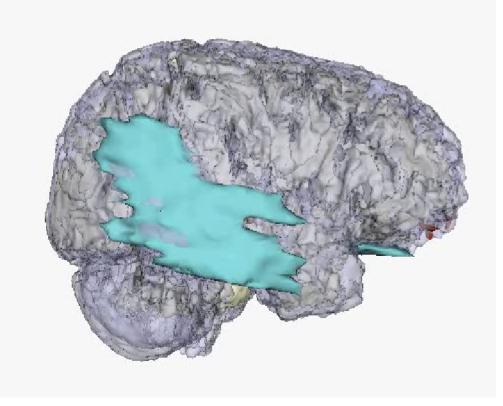
Segmentation of remaining lesions and hemorrhages in this subject can be accomplished using the tools and workflows demonstrated in this tutorial. Sample images of this undertaking are illustrated below.





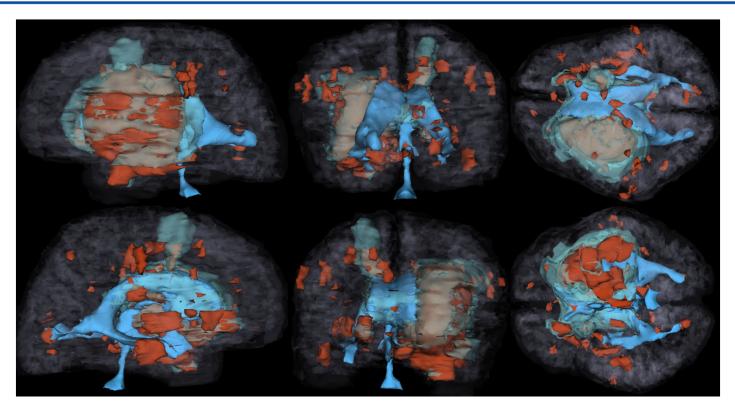






Click on image to play





Download movie of this subject's segmentation from link below:

http://wiki.na-mic.org/Wiki/index.php/File:TBISegmentation-Subject2.avi



- 3D Slicer offers powerful methodologies for the visualization of pathology due to traumatic brain injury
- Use of multiple MR image channels greatly enhances the ability to study and understand TBI structure/extent
- ABC is a robust algorithm to perform joint coregistration and automatic segmentation of TBI
- Completion of this tutorial allows one to acquire useful expertise on how to identify and characterize TBI
- Use of 3D Slicer can offer informed strategies for quantification of TBI-related edema or hemorrhage and for improved insight of clinical relevance



The members of the UCLA TBI team would like to convey our special gratitude to Dr. Ron Kikinis for his excellent support, direction and leadership of the concerted efforts that have made this work possible.

We are also indebted to Dr. Sonia Pujol for her expertise and assistance, as well as for providing us with her guidance throughout the research study that led to the creation of this tutorial.

The experience and mentorship of Drs. Randy Gollub and Steve Pieper was essential throughout the work that led to this tutorial.



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## National Institutes of Health

National Institute of Neurological Disorders and Stroke

INAL INSTITUTE OF UROLOGICAL NIH P01NS058489 (PI: Paul Vespa MD, FAAN, FCCN)



### Laboratory of Neuro Imaging

University of California, Los Angeles

Director: Arthur W. Toga PhD



- A Irimia, MC Chambers, JR Alger, M Filippou, MW Prastawa, B Wang, DA Hovda, G Gerig, AW Toga, R Kikinis, PM Vespa, JD van Horn (2011) Comparison of acute and chronic traumatic brain injury using semi-automatic multimodal segmentation of MR volumes *submitted*
- A Irimia, MC Chambers, M Filippou, JR Alger, MW Prastawa, B Wang, S Gouttard, SMA Pujol, SR Aylward, DA Hovda, G Gerig, AW Toga, R Kikinis, PM Vespa, JD van Horn (2011) Threedimensional calculation and quantification of morphometric and volumetric cortical atrophy indices of widespread clinical use from MRI volumes of traumatic brain injury using 3D Slicer *Proceedings of the 41<sup>st</sup> Annual Meeting of the Society for Neuroscience (SfN 2011), Washington, DC (this work was honored with the 1<sup>st</sup> Prize in the Fine Science Tools Research Contest of the Brain Research Institute at UCLA)*
- A Irimia, JD van Horn, MC Chambers, MW Prastawa, S Gouttard, PM Vespa, DA Hovda, JR Algers, SMA Pujol, G Gerig, SR Aylward, AW Toga, R Kikinis (2011) Automatic multimodal MR image segmentation for the clinical assessment of traumatic brain injury in 3D Slicer *Proceedings of the 17<sup>th</sup> Annual Meeting of the Organization on Human Brain Mapping (OHBM 2011), Quebec City, Canada*



#### On TBI:

Langlois, J.A., Rutland-Brown, W. and Thomas, K. (2006). Traumatic brain injury in the United States: emergency department visits, hospitalizations, and deaths.

Chen, A.J. and D'Esposito, M. (2010). Traumatic brain injury: from bench to bedside to society. Neuron. 66, 11-14.

Faul, M., Xu, L., Wald, M.M. and Coronado, V.G. (2010). Traumatic Brain Injury in the United States: Emergency Department Visits, Hospitalizations and Deaths 2002-2006.

#### On ABC:

Prastawa, M., Bullitt, E. and Gerig, G. (2009). Simulation of brain tumors in MR images for evaluation of segmentation efficacy. Med Image Anal. 13, 297-311.

Prastawa, M., Bullitt, E., Ho, S. and Gerig, G. (2004). A brain tumor segmentation framework based on outlier detection. Med Image Anal. 8, 275-283.

Prastawa, M., Bullitt, E., Moon, N., Van Leemput, K. and Gerig, G. (2003). Automatic brain tumor segmentation by subject specific modification of atlas priors. Acad Radiol. 10, 1341-1348.

Prastawa, M., Gerig, G. (2008). Brain lesion segmentation through physical model estimation. Lecture Notes in Computer Science. 5358, 562-571.

#### On lesion segmentation:

Ding, K., Marquez de la Plata, C., Wang, J.Y., Mumphrey, M., Moore, C., Harper, C., Madden, C.J., McColl, R., Whittemore, A., Devous, M.D. and Diaz-Arrastia, R. (2008). Cerebral atrophy after traumatic white matter injury: correlation with acute neuroimaging and outcome. J Neurotrauma. 25, 1433-1440.

Dubroff, J.G. and Newberg, A. (2008). Neuroimaging of traumatic brain injury. Semin Neurol. 28, 548-557.

- Greenberg, S.M., Vernooij, M.W., Cordonnier, C., Viswanathan, A., Al-Shahi Salman, R., Warach, S., Launer, L.J., Van Buchem, M.A. and Breteler, M.M. (2009). Cerebral microbleeds: a guide to detection and interpretation. Lancet Neurol. 8, 165-174.
- Huisman, T.A. (2003). Diffusion-weighted imaging: basic concepts and application in cerebral stroke and head trauma. Eur Radiol. 13, 2283-2297.
- Huisman, T.A., Sorensen, A.G., Hergan, K., Gonzalez, R.G. and Schaefer, P.W. (2003). Diffusion-weighted imaging for the evaluation of diffuse axonal injury in closed head injury. J Comput Assist Tomogr. 27, 5-11.
- Itti, L., Chang, L. and Ernst, T. (2001). Segmentation of progressive multifocal leukoencephalopathy lesions in fluid-attenuated inversion recovery magnetic resonance imaging. J Neuroimaging. 11, 412-417.

Lee, B. and Newberg, A. (2005). Neuroimaging in traumatic brain imaging. NeuroRx. 2, 372-383.