



NA-MIC

National Alliance for Medical Image Computing

<http://www.na-mic.org>

Non-rigid Registration of MR and CT images for CT-guided liver ablation

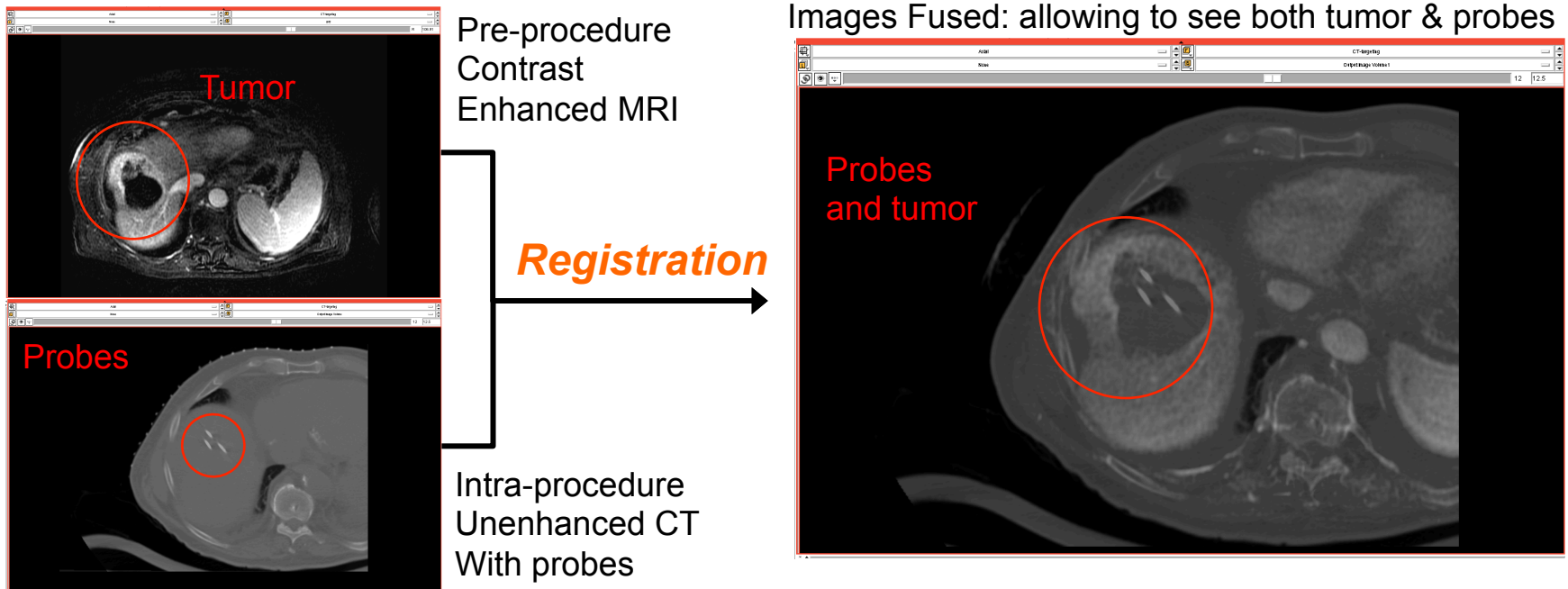
Soichiro Tani, Atsushi Yamada, Junichi Tokuda,
Dominik S. Meier, and Nobuhiko Hata

Department of Radiology
Brigham and Women's Hospital



Learning Objective

- This tutorial demonstrates how to co-register the pre-operative contrast enhanced MR image to the intra-procedure CT image via automated non-rigid registration.
- A workflow of image-guided ablation can be understood.





Tutorial Topics / Targets

Slicer Modules Used:

- N4ITK MRI Bias Correction
- Editor
- General Registration (BRAINS)

Image Processing Tasks Performed:

- Intensity non-uniformity correction
- Mask generation / segmentation
- Inter-modality non-rigid/affine image registration

Image Data Used:

- Pre-operative abdominal MR with surface coil, showing liver
- Planning CT and Guidance CT with probes



Prerequisites

- This tutorial was developed and tested on
- Windows 7, OS X 10.10.2, Ubuntu Linux 14.04
- Slicer 4.4.0-2015-01-26

This tutorial website is at:

<http://www.slicer.org/slicerWiki/index.php/Documentation/4.4/Training>

Anonymized image dataset for this tutorial is available at:

<http://www.slicer.org/slicerWiki/images/0/04/NRR-CTgLiverAblation.zip>



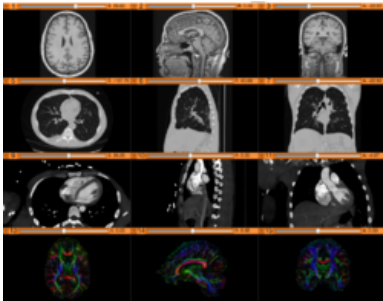
Disclaimer

Slicer is not FDA approved or CE marked and is for clinical research only.

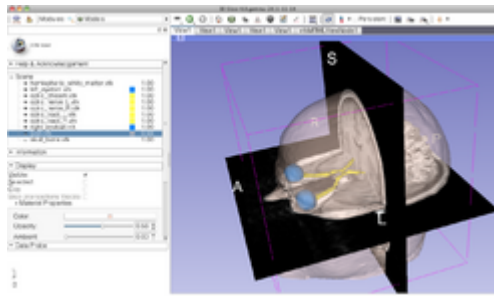


User Skill Pre-requisites

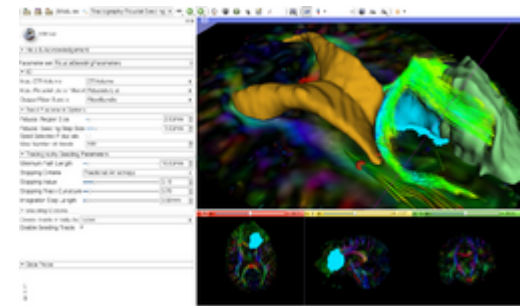
- Before running this tutorial, if new and unfamiliar with the Slicer user interface, we recommend to first complete the following tutorials, available here:
- <http://www.slicer.org/slicerWiki/index.php/Documentation/UserTraining>



[SlicerWelcome tutorial](#)



[Slicer4Minute tutorial](#)



[Neurosurgical Planning tutorial](#)

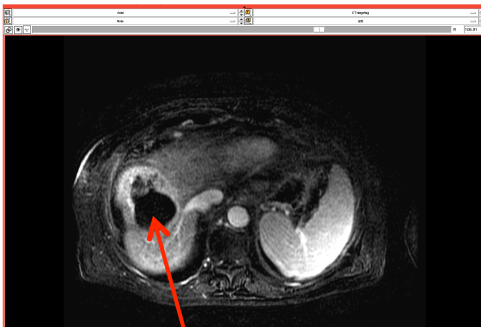


Clinical Significance

CT imaging can be used to plan an interventional approach to facilitate the safe placement of **the ablation probes** in the tumor. **However, the tumor is poorly visible on the intra-operative CT.**

Pre-operative MRI

Tumor margins and surrounding structure are visible



MRI

Tumor

Liver position, shape and structures may differ significantly between the two exams.

A non-rigid registration is desirable to compensate for liver deformation caused by patient positioning, respiratory motion and interventional manipulation.

Intra-operative CT

Only interventional probes are visible, not the tumor



Probes

CT

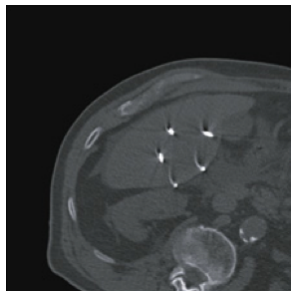


Overview of Registration Workflow

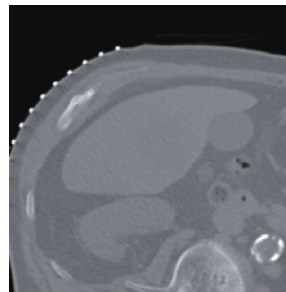
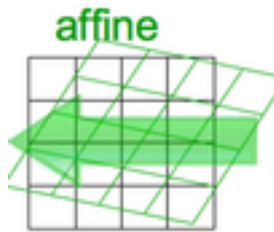
In this particular CT-guided ablation, there are two registrations necessary:

- 1) Non-rigid registration between the pre-op MR and Planning CT
- 2) Affine registration between the Planning CT and Guidance CT

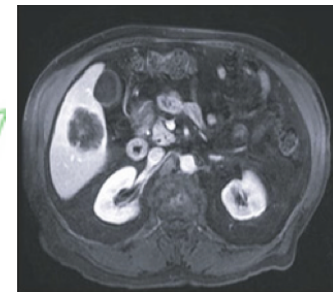
This minimizes differences between image pairs and increases robustness of the registration.



Guidance CT



Planning CT

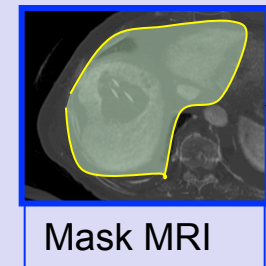
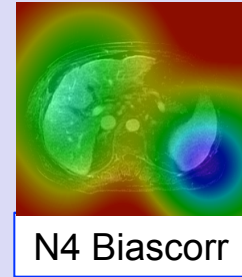
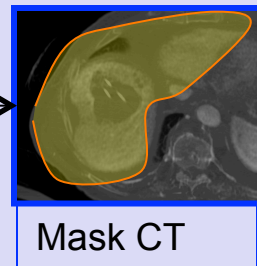


**Pre-procedure
enhanced MRI**

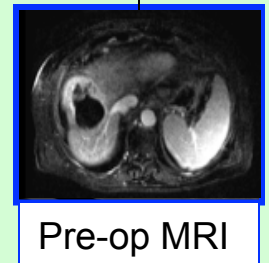
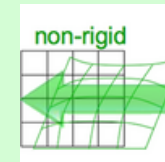
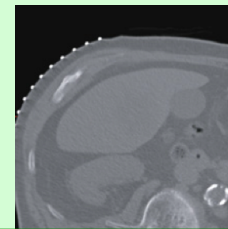
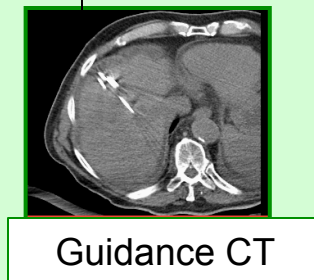


Registration Strategy/Roadmap

Preprocessing



Registration





Clinical Scenario

One month before the ablation procedure

A patient with a **possible liver tumor** is enrolled in the study. CT-guided percutaneous cryoablation is scheduled.

One week before the ablation procedure

Pre-operative contrast-enhanced MRI is acquired ("MR")

During the procedure

0 min: The patient is moved into the room

5 min: Preparation

20 min: Acquisition of planning CT that covers the entire liver ("Planning CT")

25 min: Image review for planning

40 min: Placement of probes under CT fluoroscopy guidance

45 min: Confirmation of probe placement ("Guidance CT")

60 min: Cryoablation under CT guidance



Overview of Registration Workflow

- 1. Before operation
 - A. Mask the liver on MR
 - B. Apply MRI Bias Field Intensity Correction
- 2. During operation (before insertion)
 - A. Mask the liver on Planning CT
 - B. Apply MRI-CT non-rigid registration
- 3. During operation (after insertion)
 - A. Mask the liver on Guidance CT
 - B. Apply CT-CT affine registration
 - C. Apply Resampling & Fusion

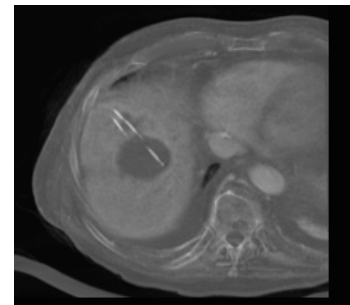
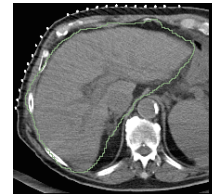
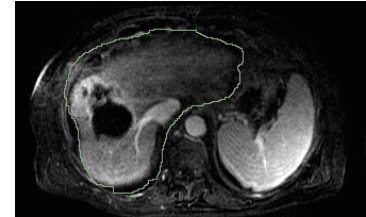
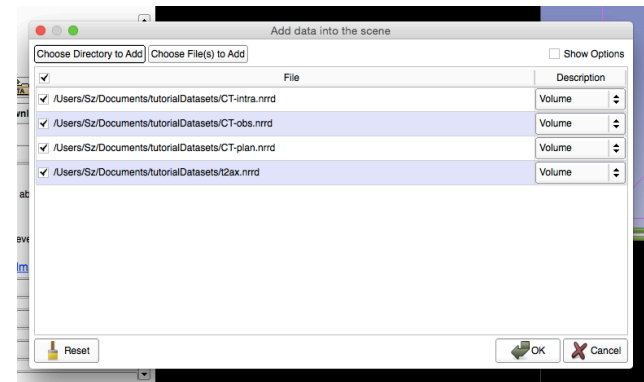
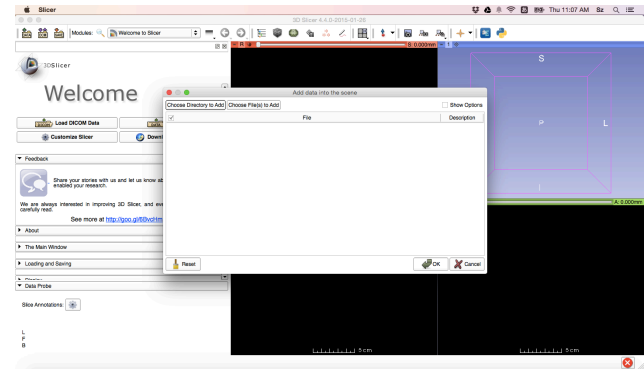




Image loading

1. Click “Load Data” on Welcome column (or “DATA” icon on toolbar)
2. “Choose Directory to Add” and select the downloaded data named “tutorial Datasets.”
3. Push “OK” after confirming that all four check boxes are checked.
4. Now all image sets are downloaded to be processed.



In real clinical cases, image importing is required in each time new images are scanned.
In this tutorial all images are downloaded at a time to avoid complexity.



1. Processing before operation

This process is done using MR images which are taken before operation.



1-A. MR mask of liver

Output: MR-label.nrrd (made automatically when Editor works)

1. Show menu and confirm image data set

First, confirm that the current background image set is “MR” which is the contrast-enhanced MR taken before operation.

The screenshot shows the Medical Image Computing Editor interface. At the top, there is a status bar with 'R' and 'S: -0.332mm'. Below it is a toolbar with various icons. A red arrow points to the 'pin icon' (a small square with a cross) in the toolbar, with the text '1. To show menu, locate your cursor to pin icon and click "<<"'. Below the toolbar is a menu with three items: 'None', 'None', and 'MR'. The 'MR' item is highlighted with a red box, and a red arrow points to it with the text '2. Select "MR" as background image'. Below the menu is a large image of a liver MRI scan. At the bottom left of the image, it says 'B: MR'. At the bottom right, there is a scale bar labeled '10 cm'.

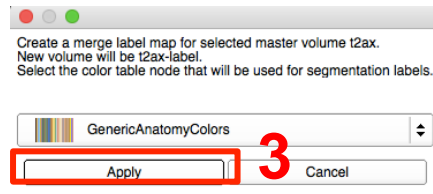
1. To show menu, locate your cursor to pin icon and click "<<"

2. Select "MR" as background image

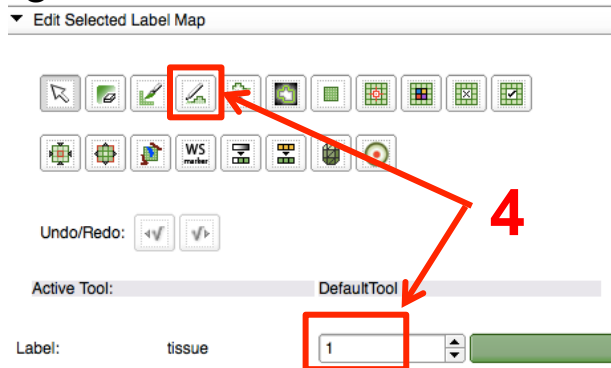


1-A. MR mask of liver

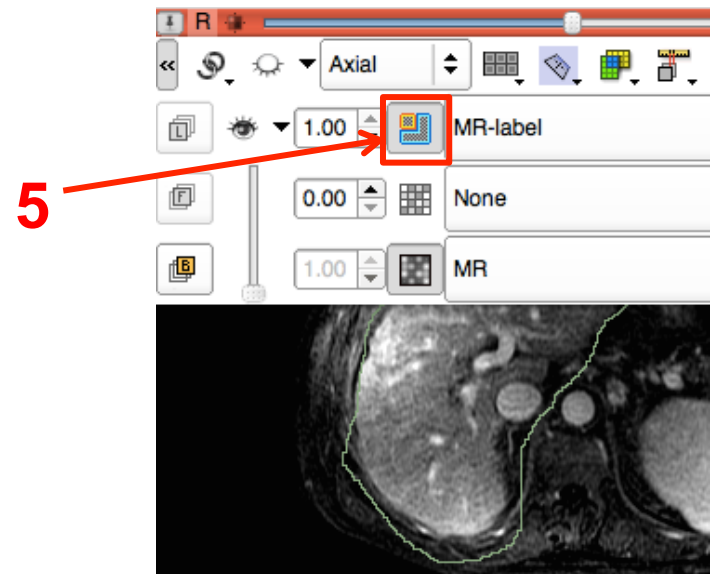
2. Go to the “Editor” module
3. Click “Apply” on the small window about Color table. This will automatically generate a new label volume, called “t2ax-label”.



4. Select “Draw” button of the module pane to segment liver with label 1



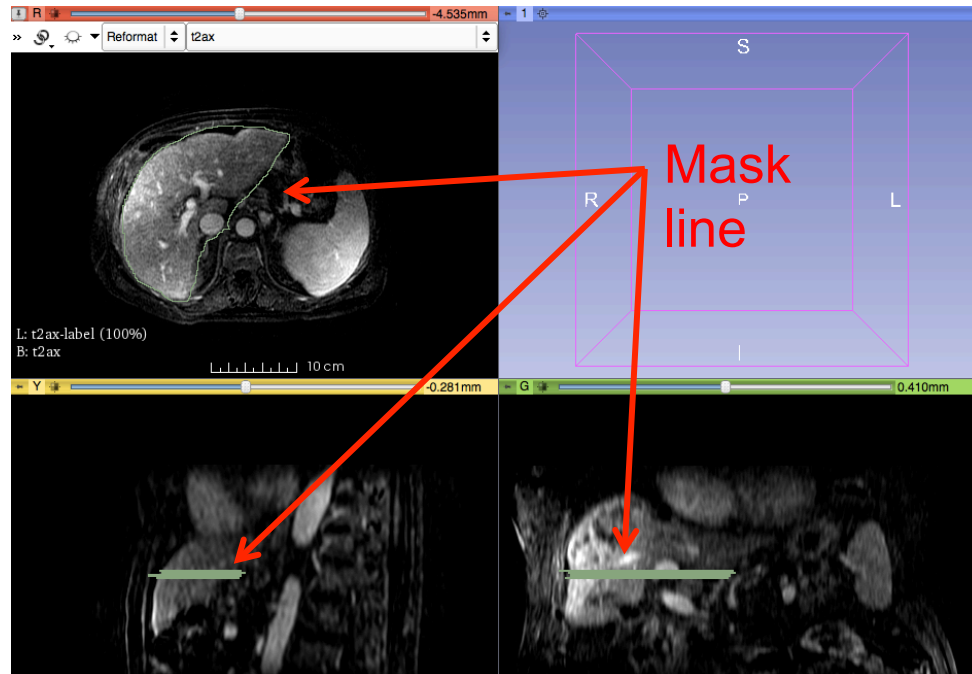
5. Choose the outlined icon below to confirm the segmented area easily





1-A. MR mask of liver

6. Draw an outer border around the liver on the axial slice. When done hit the “Return” key or “Apply” button to close the contour. Then hit arrow keys to move to the next slice.



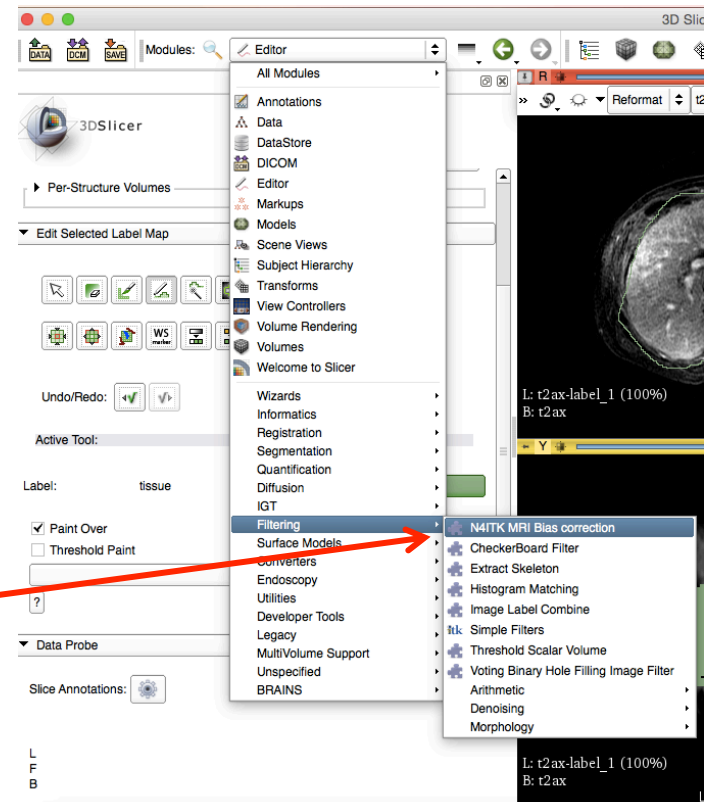


1-B. MR Bias Correction

Input: MR.nrrd and MR-label.nrrd, Output: MR-N4.nrrd

This “MR” images are obtained with surface coils that exhibit a strong intensity falloff, visible in the MR image as areas away from the surface being significantly darker. Because this can negatively affect registration quality, we correct it first. This process of intensity normalization is often called “**Bias Correction**”

Module Used: “**N4ITK MR Bias Correction**”
(in the module menu under “Filtering”)





1-B. MR Bias Correction

1. Set Parameters:

Input image = **MR**

Mask image = **MR-label**

Output volume = choose “Create and rename new Volume” and name it “**MR-N4**”

BSpline grid resolution = **4,4,4**

2. Click “Apply”*

3. Save your current work

*This process will take 1-3 minutes.

N4ITK MRI Bias correction

Parameter set: N4ITK MRI Bias correction

IO

Input Image: MR

Mask Image: MR-label

Output Volume: MR-N4

Output bias field image: None

N4 Parameters

BSpline grid resolution: 4,4,4

Spline distance: 0.00

Bias field Full Width at Half Maximum: 0.00

Advanced N4 Parameters

Restore Defaults AutoRun

Status: Idle

Cancel Apply



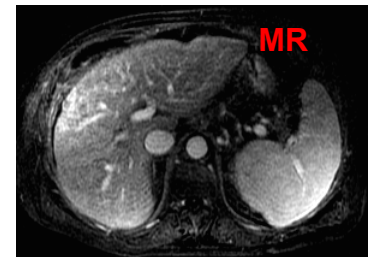
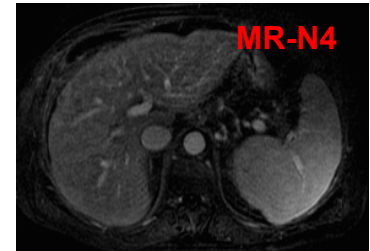
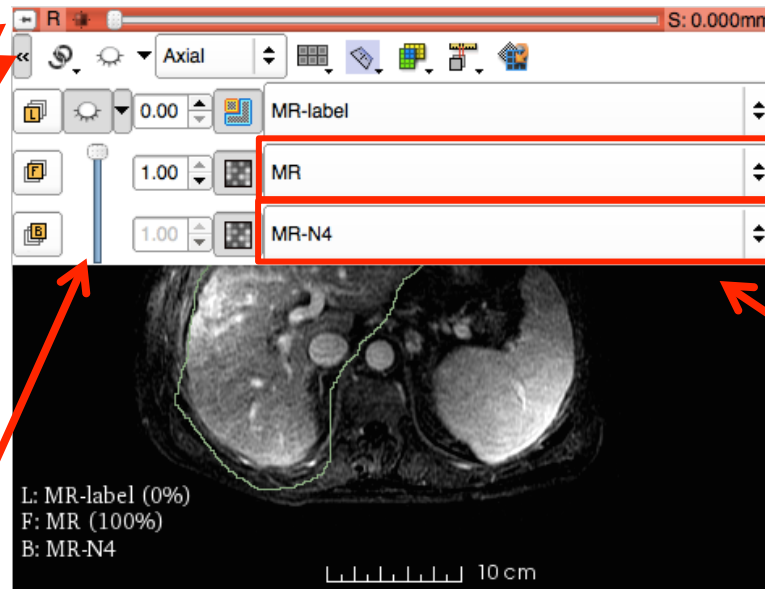
1-B. MR Bias Correction Result

Select “MR-N4” at Background layer and “MR” at Foreground layer.

Switching between background and foreground you can see the Bias Correction applied.

1. To show menu, locate your cursor to pin icon and click “<<”

3. Use the slider to fade between the background and foreground viewers



2. Set images as follows:
Foreground = MR
Background = MR-N4



2. Processing during operation (before insertion)

Now imagine that the patient is prepared for operation in a CT room.

CT scan for planning is taken before needle insertion and imported to Slicer.



2-A. Liver masking on Planning CT

Input: CTP.nrrd, Output: CTP-label.nrrd

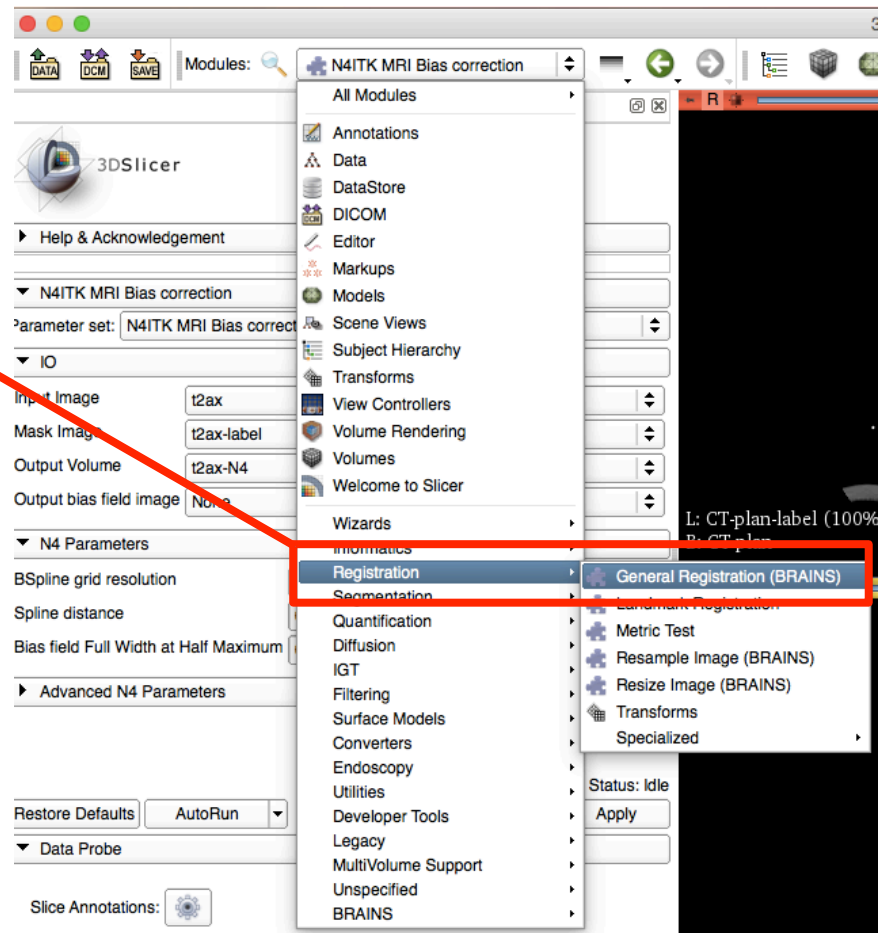
1. Save your current work (Please confirm the directory).
2. We now repeat the same outlining process on the CT image.
3. In the Editor, select under Master Volume the “CTp”.
4. As before, the editor will suggest a “CTp-label” automatically.
5. Proceed as for “1-A. MR mask of liver” above to generate a CT liver mask.
6. Again Save your work.



2-B. MRI-CT Registration

Input: CTp.nrrd, MR-N4.nrrd, CTp-label and MR-label

Go to the “General Registration (BRAINS)” module in the “Registration” category of the module menu





2-B. MRI-CT Registration

Output: T-MR-N4-to-CTp and Im-MR-N4-to-CTp

Set “General Registration (BRAINS)” module parameters as follows

- Fixed image volume = “**CTp**”
- Moving image volume = “**MR-N4**”
- B-spline Grid Size = “**5,5,5**”
- Slicer BSpline transform = “Create and rename a new transform” and name it “**T-MR-N4-to-CTp**”
- Output image volume = “Create and rename a new Volume” and name it “**Im-MR-N4-to-CTp**”
- Check “**useCenterOfROIAlign**”

▼ Input Images

Fixed Image Volume	CTp
Moving Image Volume	MR-N4
Percentage Of Samples	0.002
B-Spline Grid Size	5,5,5

▼ Output Settings (At least one output must be specified)

Slicer Linear Transform	None
Slicer BSpline Transform	T-MR-N4-to-CTp
Output Image Volume	Im-MR-N4-to-CTp

▼ Transform Initialization Settings

Initialization transform	None
Initialize Transform Mode	<input checked="" type="radio"/> useCenterOfROIAlign



2-B. MRI-CT Registration

- Select the Registration methods to run, check the following boxes:

- “Rigid (6DOF)”
- “Rigid+Scale(7DOF)”
- “Rigid+Scale+Skew(10DOF)”
- “Affine(12DOF)”
- “BSpline (>27DOF)”

- Check “**ROI**” of Mask Proceeding
- Input fixed mask = “**CTp-label**”
- Input moving mask = “**MR-label**”.

Click “Apply” and wait for 1-3 minutes.

▼ Registration Phases (Check one or more, executed in order listed)

Rigid (6 DOF)	<input checked="" type="checkbox"/>
Rigid+Scale(7 DOF)	<input checked="" type="checkbox"/>
Rigid+Scale+Skew(10 DOF)	<input checked="" type="checkbox"/>
Affine(12 DOF)	<input checked="" type="checkbox"/>
BSpline (>27 DOF)	<input checked="" type="checkbox"/>
SyN	<input type="checkbox"/>
Composite (many DOF)	<input type="checkbox"/>

▼ Image Mask and Pre-Processing

Masking Option

☐ NOMASK ☐ ROIAUTO

☒ ROI

(ROI) Masking input fixed

(ROI) Masking input moving

CTp-label

MR-label

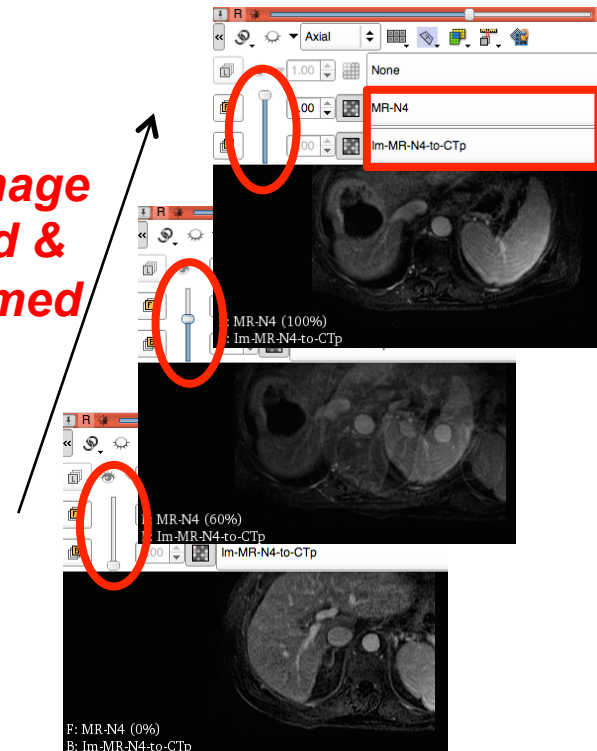
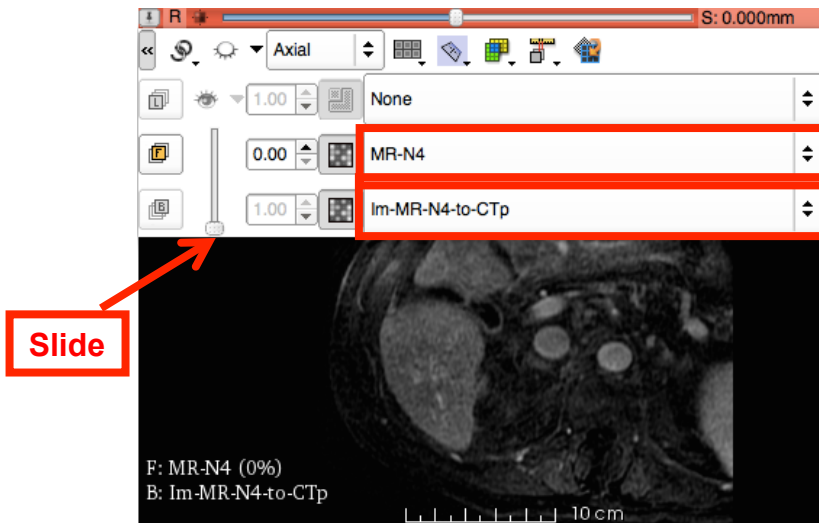


2-B. MRI-CT Registration Result – Comparison 1

Select “**Im-MR-N4-to-CTp**” at Background layer and “**MR-N4**” at Foreground layer. Switching between background and foreground you can now see the deformation applied.

Foreground: **MR-N4**
Background: Im-MR-N4-to-CTp

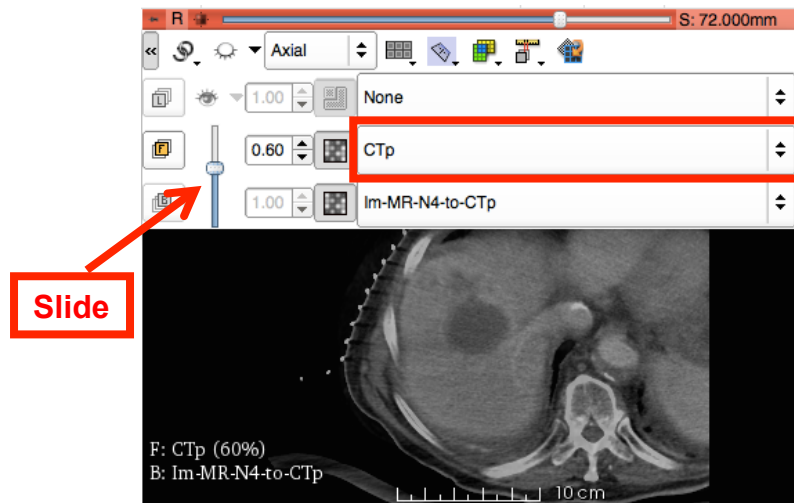
*MR image
moved &
deformed*





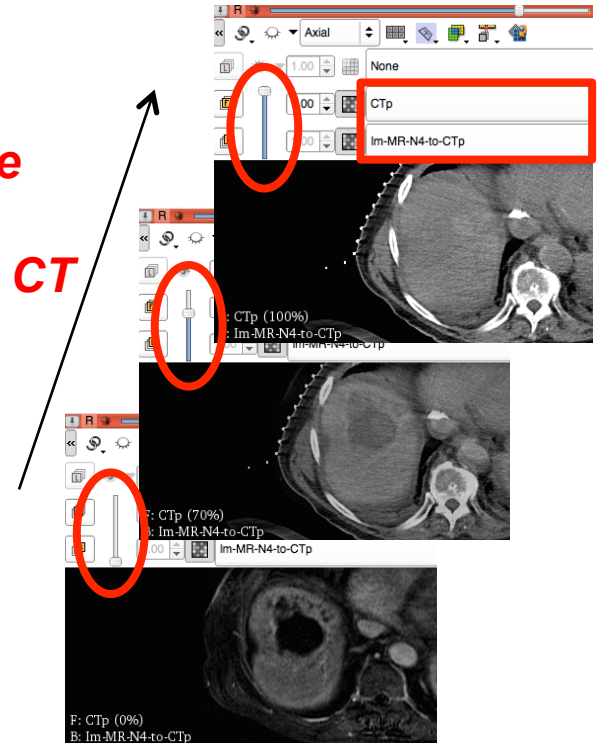
2-B. MRI-CT Registration Result – Comparison 2

Select “CTp” at Foreground layer.
You can see that the shape of the
liver on MRI was deformed and
fitted the liver on CT image.



Foreground: **CTp**
Background: Im-MR-N4-to-CTp

*MR image
fitted
Planning CT*





3. Processing during operation (after insertion)

The liver tumor is targeted and needles are inserted by the physician.

CT image for guidance is taken to confirm location of needles.



3-A. Liver masking on Guidance CT

Input: CTg.nrrd Output: CTg-label

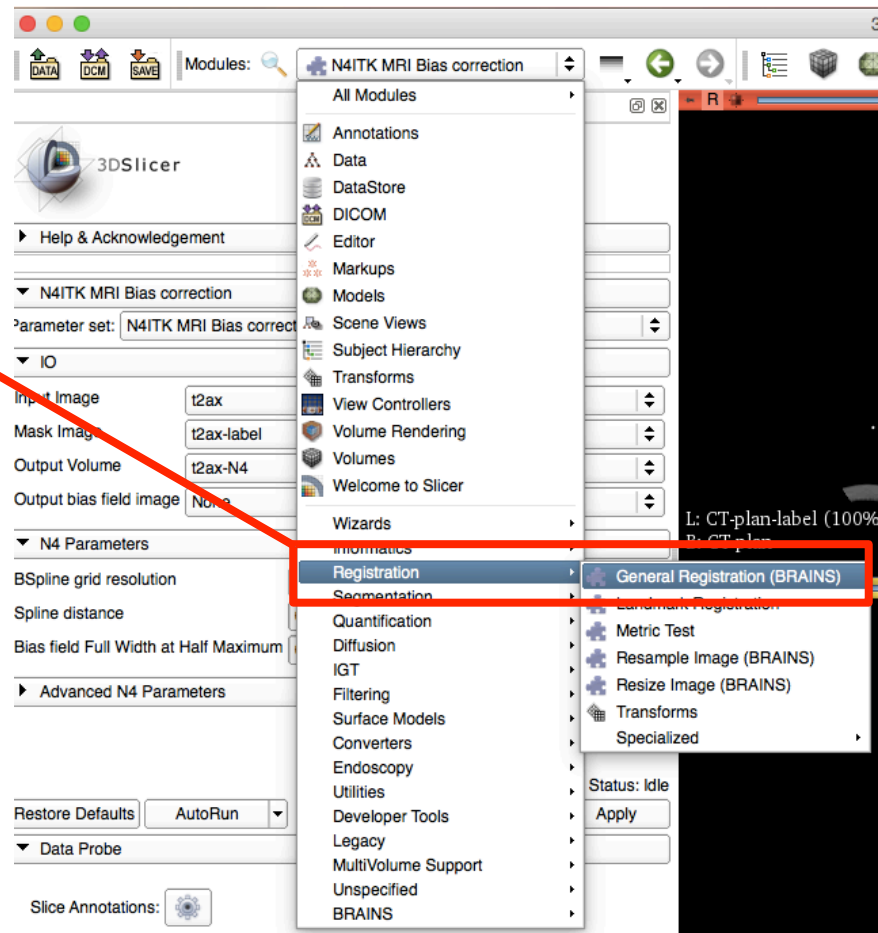
1. Save your current work.
2. We now repeat the same outlining process on the CT image.
3. In the Editor, select under Master Volume the “**CTg**”.
4. As before, the editor will automatically suggest a “**CTg-label**” automatically.
5. Proceed as for “1-A. MR mask of liver” above to generate a CT liver mask
6. Again Save your work.



3-B. Planning CT to Guidance CT Affine Registration

Input: CTg.nrrd, CTp.nrrd, CTg-label and CTp-label

Go to the “General Registration (BRAINS)” module in the “Registration” category of the module menu





3-B. Planning CT to Guidance CT Affine Registration

We apply affine registration here.

Deference between “Planning CT” and “Guidance CT” is supposed to be small because these two image sets are taken in the same operation in the same modality while the patient is kept lying down.

The obtained transform data here is to be applied at 3-C.



3-B. Planning CT to Guidance CT Affine Registration

Output: T-CTp-to-CTg, Im-CTp-to-CTg

Set “General Registration (BRAINS)” module parameters as follows.

- Fixed image volume = “**CTg**”
- Moving image volume = “**CTp**”
- Slicer Linear transform = “Create and rename a new transform” and name it “**T-CTp-to-CTg**”
- Slicer BSpline transform = “**None**”
- Output image volume = “Create and rename a new Volume” and name it “**Im-CTp-to-CTg**”

▼ Input Images	
Fixed Image Volume	CTg
Moving Image Volume	CTp
Percentage Of Samples	0.002
B-Spline Grid Size	5,5,5
▼ Output Settings (At least one output must be specified)	
Slicer Linear Transform	T-CTp-to-CTg
Slicer BSpline Transform	None
Output Image Volume	Im-CTp-to-CTg
▼ Transform Initialization Settings	
Initialization transform	None
Initialize Transform Mode	<input type="radio"/> Off <input type="radio"/> u <input type="radio"/> useCenterOfHeadAlign <input type="radio"/> u <input checked="" type="radio"/> useCenterOfROIAlign



3-B. Planning CT to Guidance

CT Affine Registration

- Uncheck “BSpline (>27DOF)” box
- Input fixed mask = “**CTg-label**”
- Input moving mask = “**CTp-label**”

Click “Apply” and you can see deformed CTp image as “**Im-CTp-to-CTg**”. Please save your work.

Registration Phases (Check one or more, executed in order listed)

Rigid (6 DOF)	<input checked="" type="checkbox"/>
Rigid+Scale(7 DOF)	<input checked="" type="checkbox"/>
Rigid+Scale+Skew(10 DOF)	<input checked="" type="checkbox"/>
Affine(12 DOF)	<input checked="" type="checkbox"/>
BSpline (>27 DOF)	<input type="checkbox"/>
SyN	<input type="checkbox"/>
Composite (many DOF)	<input type="checkbox"/>

Image Mask and Pre-Processing

Masking Option

☐ NOMASK ☐ ROIAUTO

☒ ROI

(ROI) Masking input fixed

(ROI) Masking input moving

(ROIAUTO) Output fixed mask

(ROIAUTO) Output moving mask

CTg-label

CTp-label

None

None

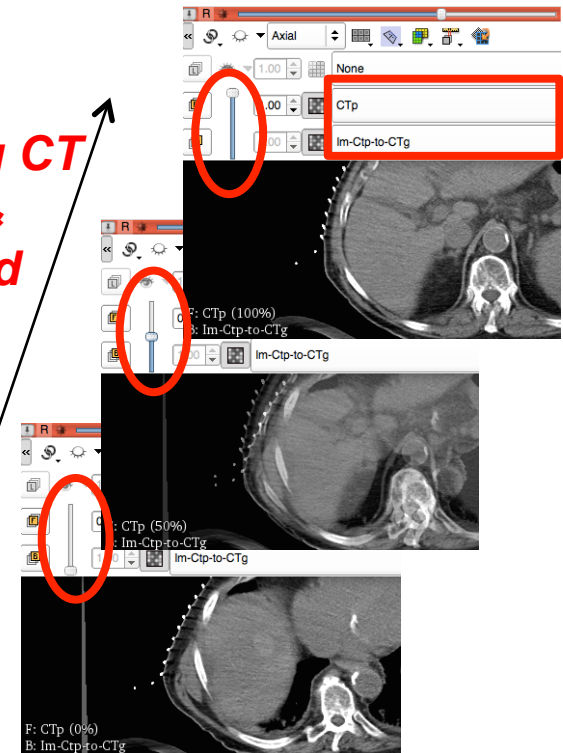
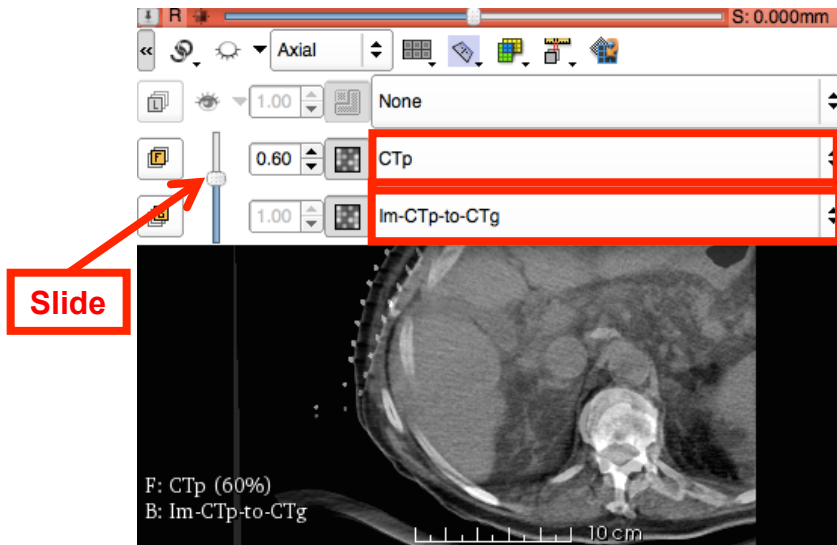


3-B. CT-CT Registration Result – Comparison 1

Select “Im-CTp-to-CTg” at Background layer and “CTp” at Foreground layer. Switching between background and foreground you can now see the deformation applied.

Foreground: **CTp**
Background: Im-CTp-to-CTg

*Planning CT
moved &
deformed*



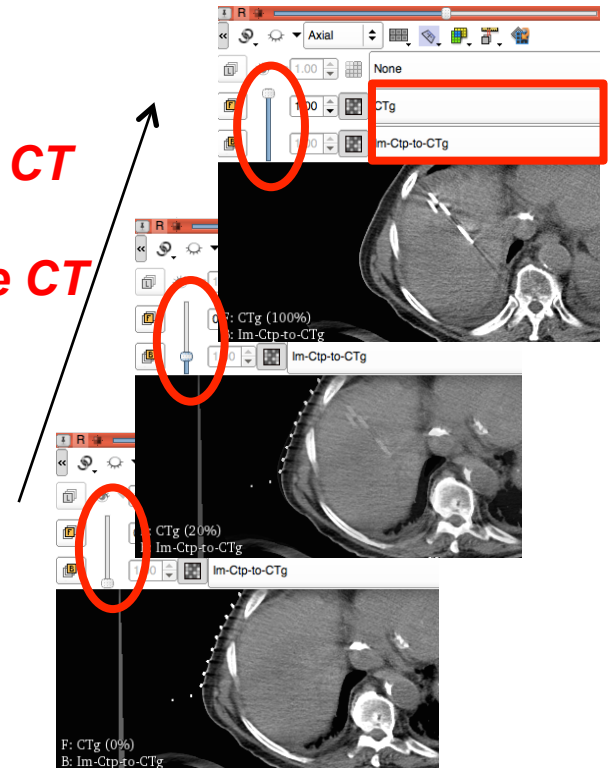
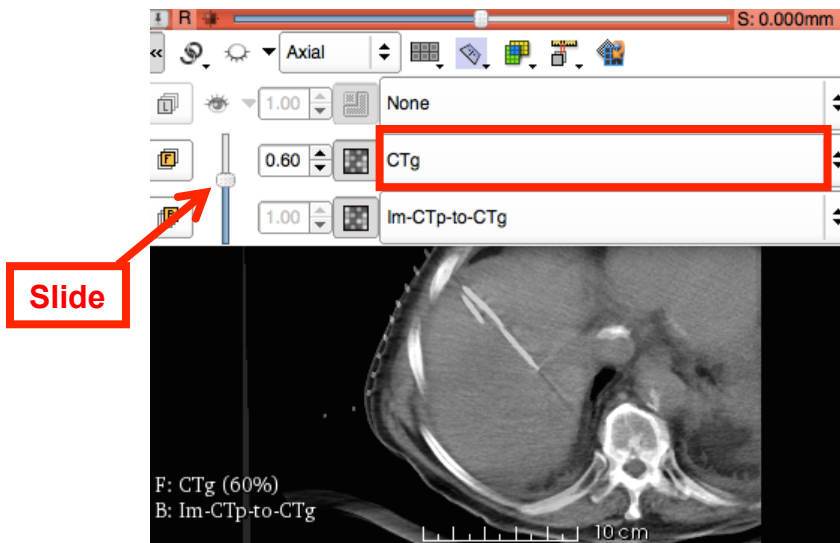


3-B. CT-CT Registration Result – Comparison 2

Select “CTg” at Foreground layer.
You can see that the shape of the
liver on CTp was deformed and
fitted the liver on CTg image.

Foreground: **CTg**
Background: Im-CTp-to-CTg

*Planning CT
fitted
Guidance CT*





3-C. Resampling and Fusion

Input: Im-MR-N4-to-CTp.nrrd, CTg, T-CTp-to-CTg Output: Im-MR-N4-to-CTp-to-CTg

Transform the deformed MR again following the CT-CT registration data

1. Go to Resample Image (Brains) module
2. Set parameters as follows:

Image To Warp = “Im-MR-N4-to-CTp”

Reference Image = “CTg”

Output Image = create and rename new Volume as “Im-MR-N4-to-CTp-to-CTg”

Transform file= “T-CTp-to-CTg”

Check Interpolation mode “Linear”

3. Click “Apply” and you can see the moved and deformed Im-MR-to-CTp image as “Im-MR-N4-to-CTp-to-CTg”.

The screenshot shows the 'Resample Image (Brains)' module interface. The 'Inputs' section has 'Image To Warp' set to 'Im-MR-N4-to-CTp' and 'Reference Image' set to 'CTg'. The 'Outputs' section has 'Output Image' set to 'Im-MR-N4-to-CTp-to-CTg'. The 'Pixel Type' section has 'float' selected. The 'Warping Parameters' section has 'Displacement Field (deprecated)' set to 'None', 'Transform file' set to 'T-CTp-to-CTg', and 'Interpolation Mode' set to 'Linear'.

Inputs	
Image To Warp	Im-MR-N4-to-CTp
Reference Image	CTg

Outputs	
Output Image	Im-MR-N4-to-CTp-to-CTg

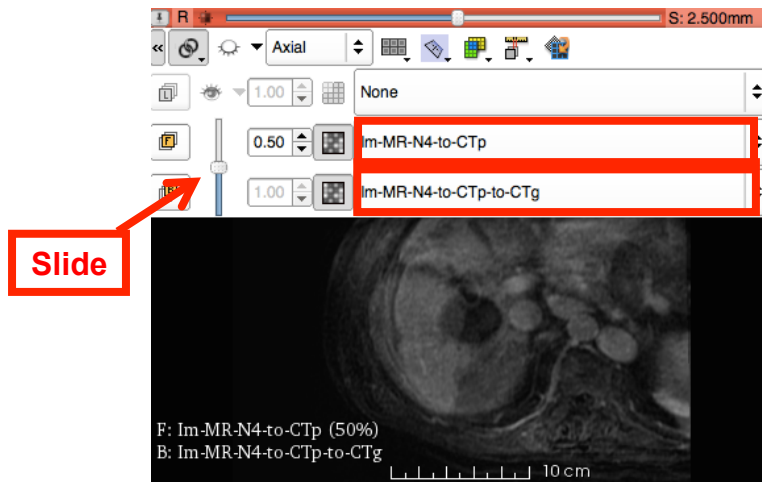
Pixel Type: ☒ float ☐ short ☐ ushort ☐ int ☐ uint ☐ u ☐ binary

Warping Parameters	
Displacement Field (deprecated)	None
Transform file	T-CTp-to-CTg
Interpolation Mode	<input checked="" type="radio"/> Linear <input type="radio"/> NearestNeighbor <input type="radio"/> ResampleInPlace <input type="radio"/> BSpline



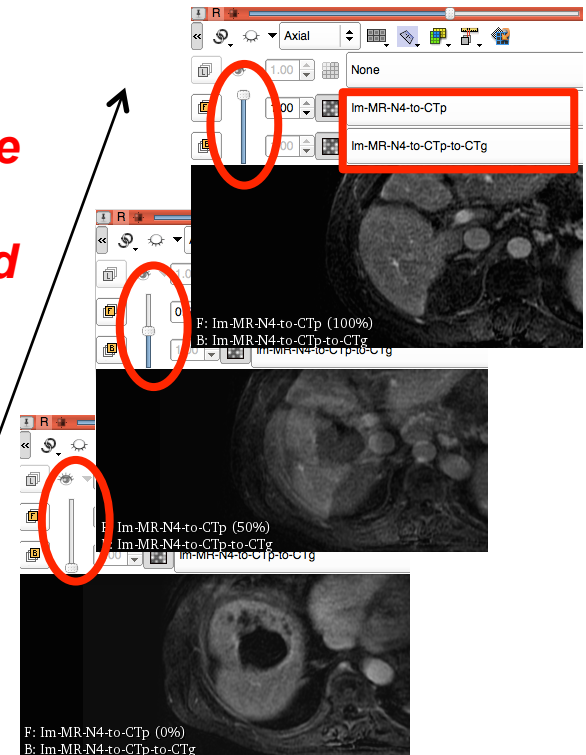
3-C. MRI-CT Resampling Result – Comparison 1

Select “**Im-MR-N4-to-CTp-to-CTg**” at Background layer and “**Im-MR-N4-to-CTp**” at Foreground layer. Switching between background and foreground you can now see the deformation applied.



Foreground: **Im-MR-N4-to-CTp**
Background: Im-MR-N4-to-CTp-to-CTg

*MR image
Moved &
Deformed
again*



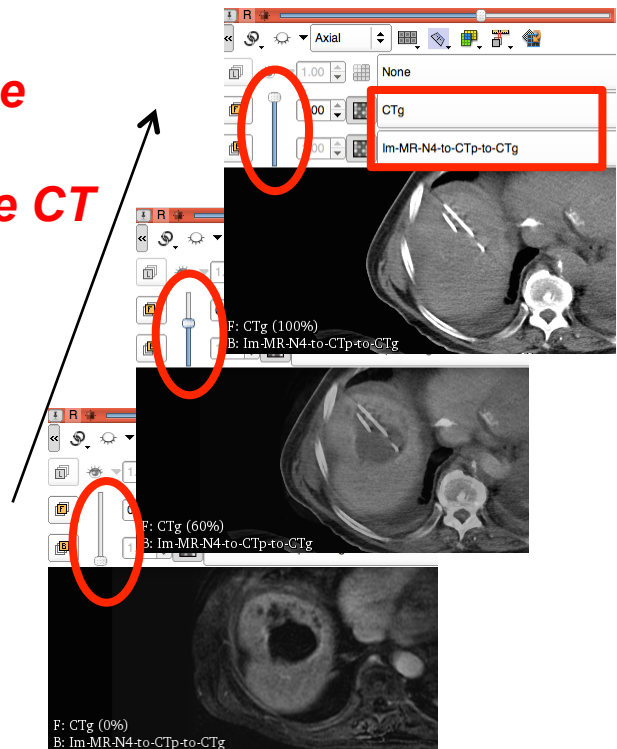
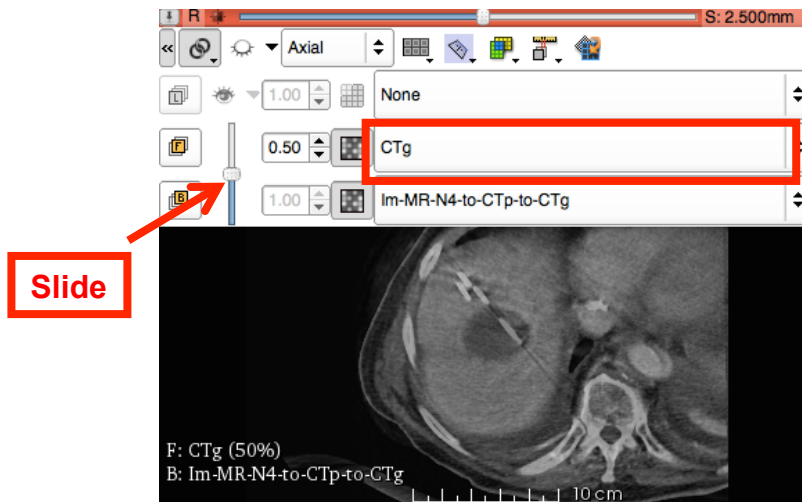


3-C. MRI-CT Resampling Result – Comparison 2

Select “**CTg**” at Foreground layer. You can see that the shape of the liver on **Im-MR-N4-to-CTp-to-CTg** is fitted the liver on CTg image.

Foreground: **CTg**
Background: Im-MR-N4-to-CTp-to-CTg

*MR image
fitted
Guidance CT*





Ablation and confirmation of margins

Now ablation is performed and CT images are taken to confirm the ablated area.

If ablated area covers the tumor with sufficient margins, the operation ends.



Conclusion

- Deformable registration and rigid registration are done on 3D slicer 4 following the process of a CT guided ablation case.
- For this particular type of image data, masking of the region of interest is usually necessary to obtain a good result.
- This registration technique can be applied to other scenario of clinical research in surgical navigation.



References

Registration workflow

- Fedorov A, Tuncali K, Fennessy FM, Tokuda J, Hata N, Wells WM, et al. Image registration for targeted MRI-guided transperineal prostate biopsy. J Magn Reson Imaging. 2012;36(4):987–92.



3D Slicer

- Gering DT, Nabavi A, Kikinis R, Hata N, O'Donnell LJ, Grimson WE, et al. An integrated visualization system for surgical planning and guidance using image fusion and an open MR. J Magn Reson Imaging. 2001;13(6):967–75.
- Fedorov A, Beichel R, Kalpathy-Cramer J, Finet J, Fillion-Robin J-CC, Pujol S, et al. 3D Slicer as an image computing platform for the Quantitative Imaging Network. Magn. Reson. Imaging. 2012 Nov;30(9):1323–41.

Bias correction

- Boyes RG, Gunter JL, Frost C, Janke AL, Yeatman T, Hill DLG, et al. Intensity non-uniformity correction using N3 on 3-T scanners with multichannel phased array coils. Neuroimage. 2008 Feb 15;39(4):1752–62.
- Tustison NJ, Avants BB, Cook PA, Zheng Y, Egan A, Yushkevich PA, et al. N4ITK: improved N3 bias correction. IEEE Trans. Med. Imaging. 2010 Jun;29(6):1310–20.



References

BRAINS Fit

- Johnson HJ, Harris G, Williams K. BRAINSFit: Mutual Information Registrations of Whole-Brain 3D Images, Using the Insight Toolkit. Insight J. 2007;
- Elhawary H, Oguro S, Tuncali K, Morrison PR, Tatli S, Shyn PB, et al. Multimodality non-rigid image registration for planning, targeting and monitoring during CT-guided percutaneous liver tumor cryoablation. Acad Radiol. Elsevier Ltd; 2010 Nov;17(11):1334–44.

B-spline registration, Mattes mutual information

- Rueckert D, Sonoda LI, Hayes C, Hill DLG, Leach MO, Hawkes DJ. Nonrigid registration using free-form deformations: application to breast MR images. Med. Imaging, IEEE Trans. 1999;18(8):712–21.
- Wells 3rd WM, Viola P, Atsumi H, Nakajima S, Kikinis R. Multi-modal volume registration by maximization of mutual information. Med Image Anal. 1996;1(1): 35–51.
- Mattes D, Haynor DR, Vesselle H, Lewellen TK, Eubank W. PET-CT image registration in the chest using free-form deformations. Med. Imaging, IEEE Trans. 2003;22(1):120–8.



Acknowledgements

Development of this tutorial is supported in part by the US National Institutes of Health (NIH) (P41 EB015898, R01 CA138586), and the Ministry of Education, Culture, Sports, Science, and Technology, Japan. The content of the material is solely the responsibility of the authors and does not necessarily represent the official views of these agencies.