

# Non-rigid Registration of Longitudinal Brain Tumor Treatment MRI

*Nicha Chitphakdithai, Veronica L. Chiang, and  
James S. Duncan*

*Image Processing and Analysis Group, Yale University  
Department of Neurosurgery, Yale School of Medicine*

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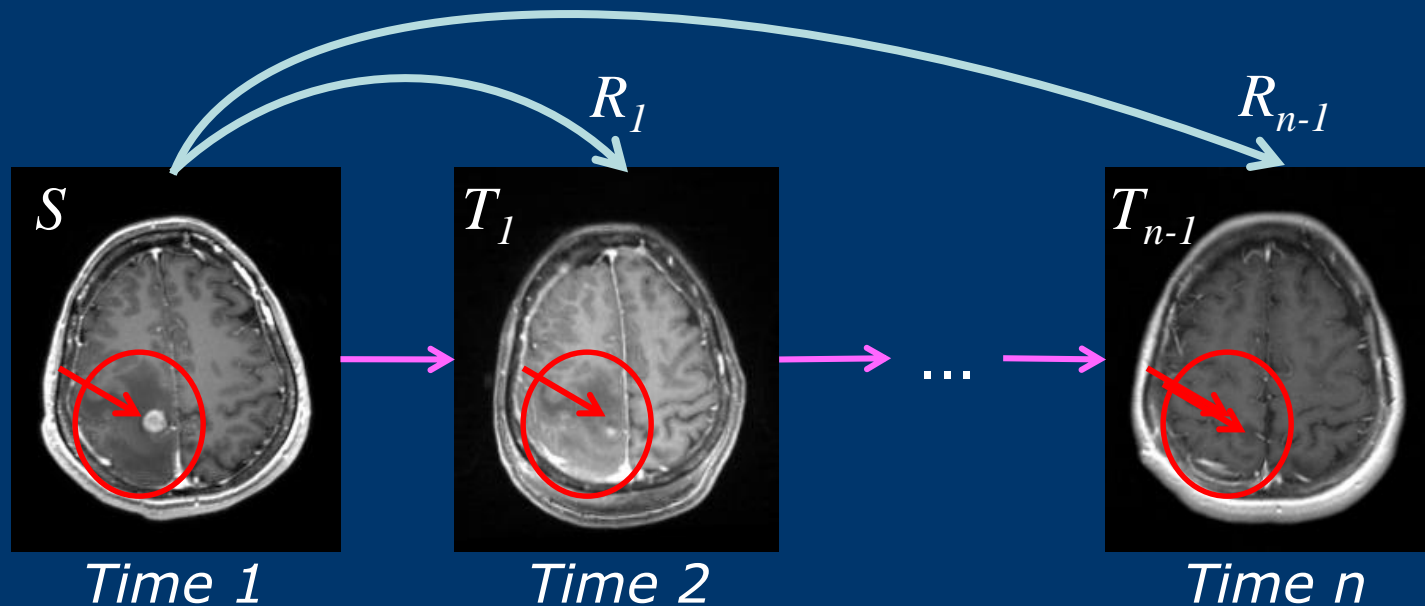
September 2, 2011



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# Image Registration Goal

- Align longitudinal brain tumor treatment MRI



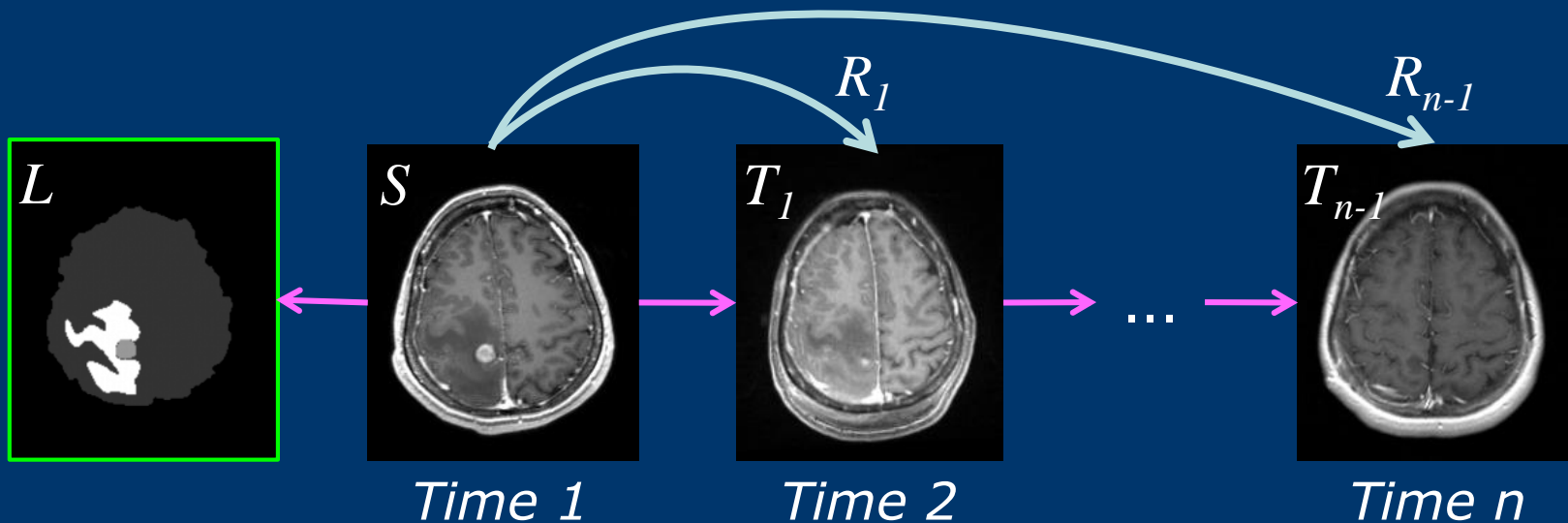
- Challenge: **Missing Correspondences**
  - Changes in tumors, edema, resection...

# Some Approaches to Handle Missing Correspondences

- First extract points for matching<sup>1</sup>
- Simulate tumor growth in normal brain<sup>2</sup>
- Adapted demons registration<sup>3</sup>
- Simultaneously estimate missing data<sup>4</sup>
  - No prior information
- Our recent work aligning brain resection MRI<sup>5</sup>
  - NOW: extend to longitudinal brain tumors, with
    - Intensity information
    - Allowing larger deformations near lesion

<sup>1</sup>Liu et al., ISBI 2010; <sup>2</sup>Zacharaki et al., Neuroimage 2009; <sup>3</sup>Risholm et al., IPMI 2009  
<sup>4</sup>Periaswamy and Farid, MedIA 2006; <sup>5</sup>Chitphakdithai and Duncan, MICCAI 2010

# Registration Framework: Setup



- Include “hidden” label map  $L$  to indicate matching vs. missing correspondences
- Maximum a posteriori framework:

$$\hat{R} = \arg \max_R \log \sum_L p(R, L | S, T)$$

# Registration Framework: EM Algorithm

- E-Step: Label Map Weights  $p(L(\mathbf{x})=l | S, T, R^k) =$   
*Likelihood*
*Intensity Prior*
*Label Map Prior*

$$\frac{\overbrace{p(T(R^k(\mathbf{x})) | S, R^k, L(\mathbf{x})=l)}^{\text{Likelihood}} \overbrace{p(S(\mathbf{x}) | L(\mathbf{x})=l)}^{\text{Intensity Prior}} \overbrace{p(L(\mathbf{x})=l)}^{\text{Label Map Prior}}}{\sum_{l'} p(T(R^k(\mathbf{x})) | S, R^k, L(\mathbf{x})=l') p(S(\mathbf{x}) | L(\mathbf{x})=l') p(L(\mathbf{x})=l')}$$

- M-Step: Update registration  $R^{k+1} =$

$$\arg \max_R \left[ \sum_{\mathbf{x} \in S} \sum_{l \in L} \overbrace{p(L(\mathbf{x})=l | S, T, R^k)}^{\text{E-Step Weights}} \log \overbrace{p(T(R(\mathbf{x})) | S, R, L(\mathbf{x})=l)}^{\text{Likelihood}} \right. \\ \left. + \sum_{\mathbf{x} \in S} \log \underbrace{p(R(\mathbf{x}) | L^{k+1})}_{\text{Transformation Prior}} \right]$$

# Probability Models: Likelihood

- Likelihood acts like similarity measure

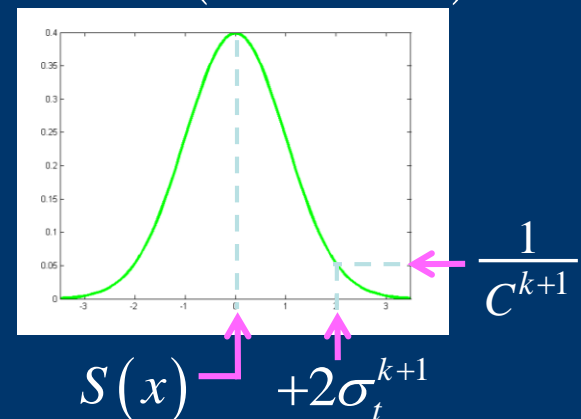
$$p(T(R(\mathbf{x})) | S, R, L(\mathbf{x}) = l) \sim \begin{cases} N(S(\mathbf{x}), \sigma_t) & , l = \text{valid tissue} \\ N(S(\mathbf{x}), \sigma_b) & , l = \text{background} \\ \text{Unif}(\frac{1}{C}) & , l = \text{abnormal class} \end{cases}$$

- Automatically update  $\sigma_t$  and  $C$ :

$$\sigma_t^{k+1} =$$

$$\sqrt{\frac{\sum_{x \in S} p(L(\mathbf{x}) = l | S, T, R^k) (T(R^k(\mathbf{x})) - S(\mathbf{x}))^2}{\sum_{x \in S} p(L(\mathbf{x}) = l | S, T, R^k)}}$$

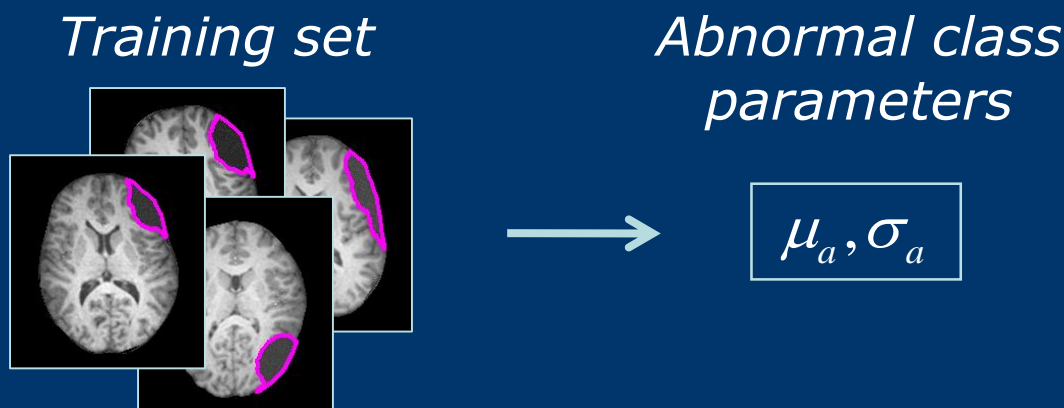
*Normal*( $S(x), \sigma_t^{k+1}$ )



# Probability Models: Intensity Prior

- Assume MRI signal characterized by tissue type  
→ Different distribution for intensities given label

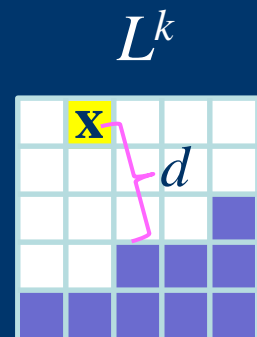
$$p(S(\mathbf{x}) | L(\mathbf{x}) = l) \sim \begin{cases} \text{Unif}\left(\frac{1}{K}\right) & , l = \text{valid tissue} \\ N(0, \sigma_b) & , l = \text{background} \\ N(\mu_a, \sigma_a) & , l = \text{abnormal class} \end{cases}$$



# Probability Models: Transformation Prior

- Transformation Model: FFDs based on uniform cubic B-splines
- Want to tolerate greater deformation near “abnormal regions” in current map estimate  $L^k$
- Model  $R(\mathbf{x}) | L^k \sim N(\mu_{\mathbf{x}}, \sigma_{\mathbf{x}})$ 
  - $\mu_{\mathbf{x}}$  = Position of voxel  $\mathbf{x}$
  - Increased  $d \rightarrow$  Decreased  $\sigma_{\mathbf{x}}$

$$\sigma_{\mathbf{x}}^2 = \begin{cases} \sigma_{\min}^2 + \frac{d_{\text{tol}} - d}{d_{\text{tol}}} (\sigma_{\max}^2 - \sigma_{\min}^2) & , d \leq d_{\text{tol}} \\ \sigma_{\min}^2 & , d > d_{\text{tol}} \end{cases}$$



■ = Abnormal Class

- Variances updated every M-step

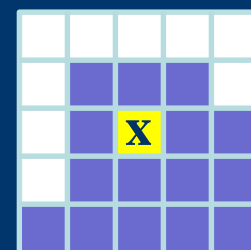


# Probability Models: Label Map Prior

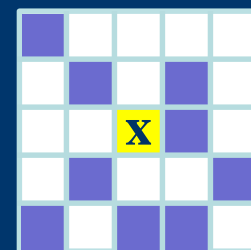
- Impose Markov random field onto label map
- Use mean-field approximation and Potts smoothing model:

$$p(L(\mathbf{x})=l) \approx \frac{1}{Z'_{\mathbf{x}}(\beta)} \exp \left[ \beta \sum_{\mathbf{n} \in N(\mathbf{x})} \delta(l, L^k(\mathbf{n})) \right],$$

$$L^k(\mathbf{n}) = \arg \max p(L(\mathbf{x})=l | S, T, R^{k-1})$$



*Higher*  
 $p(L(\mathbf{x}) = \blacksquare)$



*Lower*  
 $p(L(\mathbf{x}) = \blacksquare)$

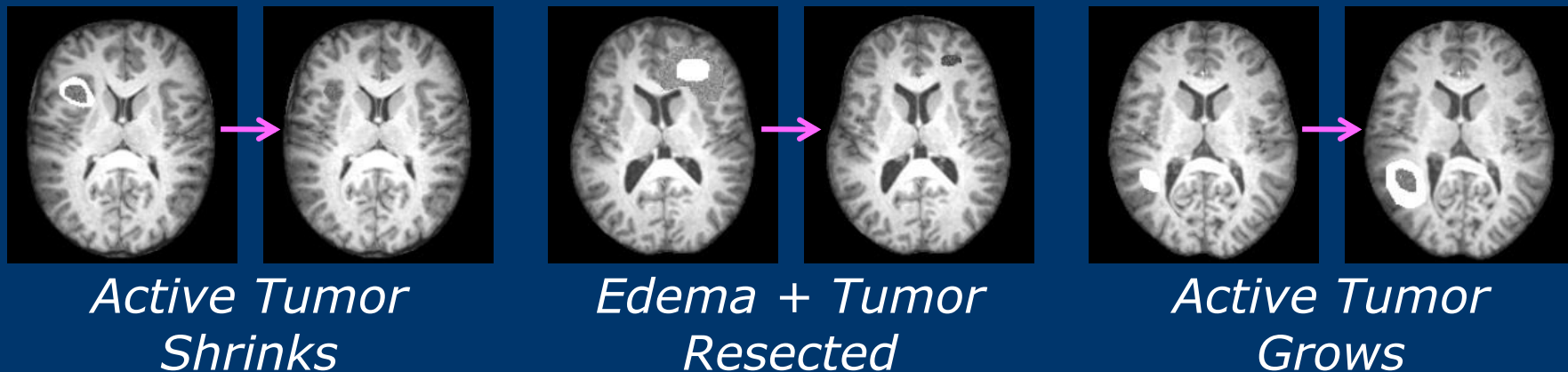
- Update  $\beta^{k+1}$  every M-step

$$\arg \max_{\beta} \sum_{\mathbf{x} \in S} \sum_{l \in L} p(L(\mathbf{x})=l | S, T, R^k) \left[ \log p(L(\mathbf{x})=l | \beta) + \log p(\beta) \right]$$

*Normal with small  $\mu, \sigma$*  ←

# Synthetic Examples: Data and Experiments

- 3 Sample Series:

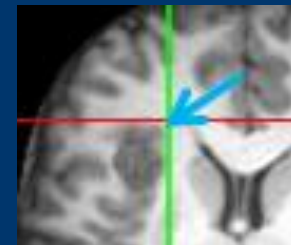
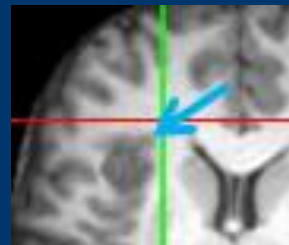
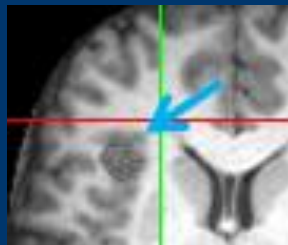
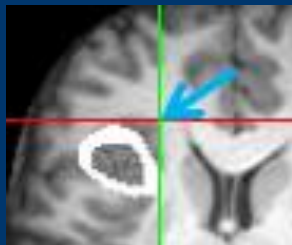


- Compared our method (*RALE*) to “standard” non-rigid registration<sup>1</sup> (*SNRR*)
  - \* Both use:
    - Voxel intensity-based similarity
    - FFD transformation model

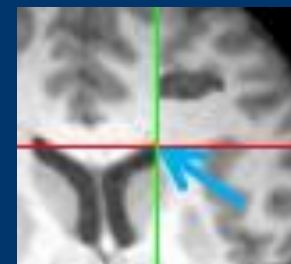
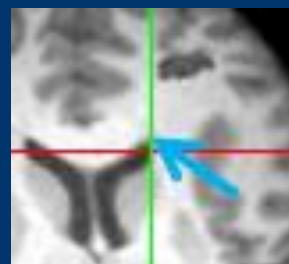
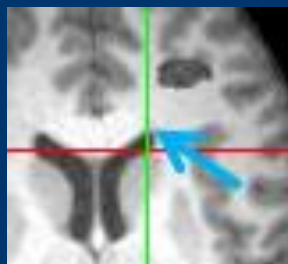
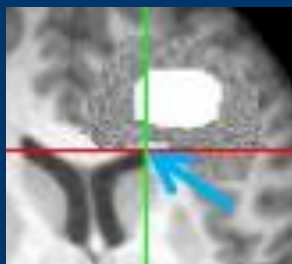
<sup>1</sup>Rueckert et al., TMI 1999

# Synthetic Examples: Sample Results

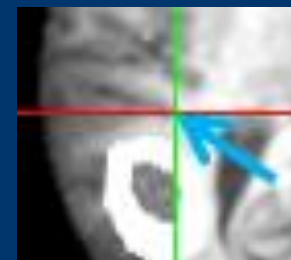
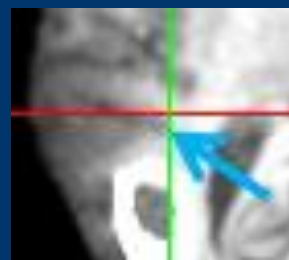
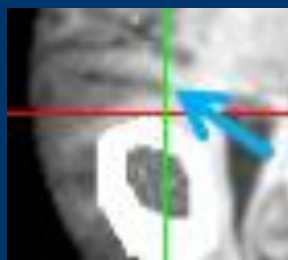
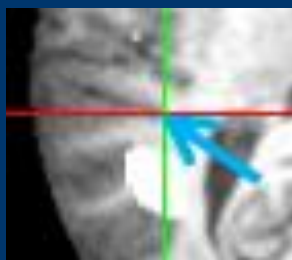
*Tumor  
Shrinks*



*Tumor  
Resected*



*Tumor  
Grows*



*Time 1*

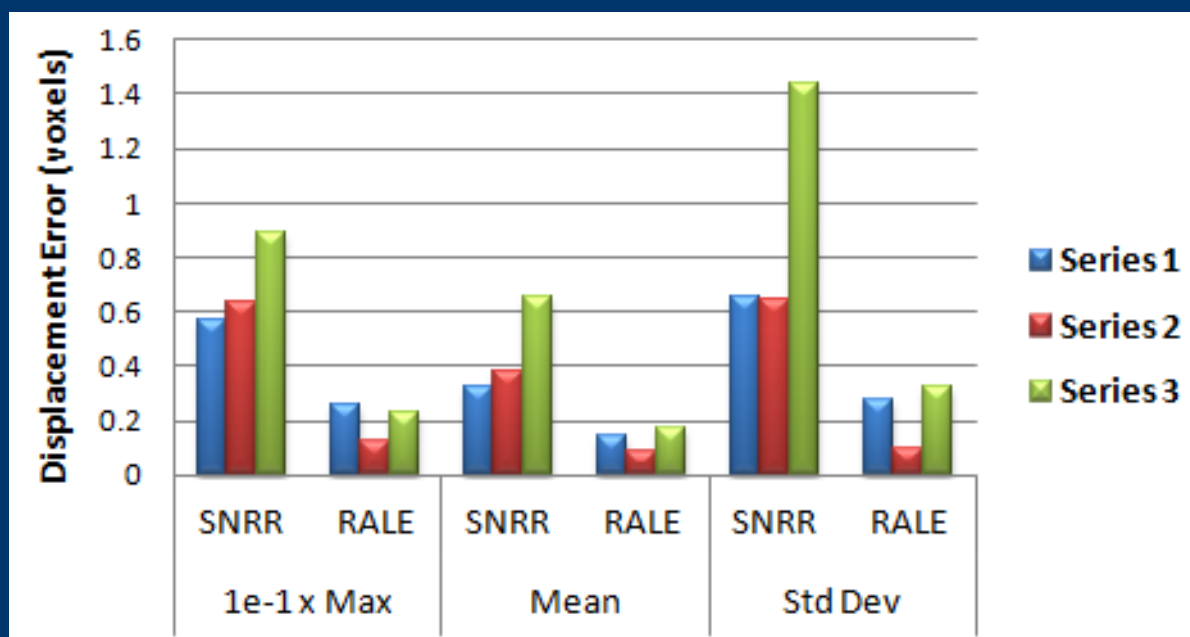
*Time 2*

*SNRR*

*RALE*

# Synthetic Examples: Registration Errors

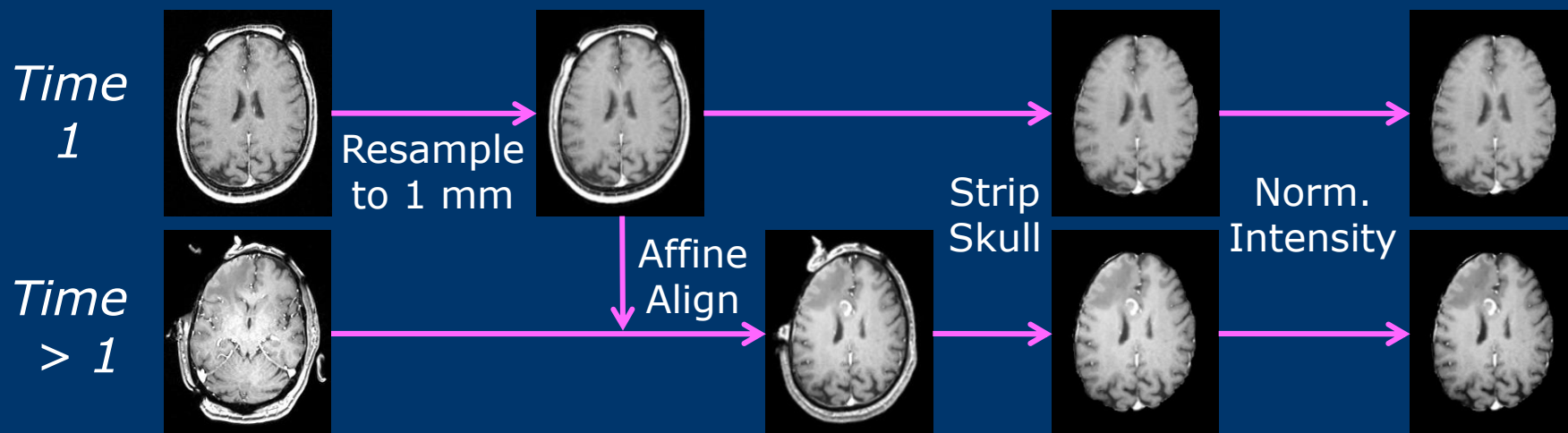
## Average Displacement Field Errors



RALE significantly reduced each error measure ( $p < 0.05$ )

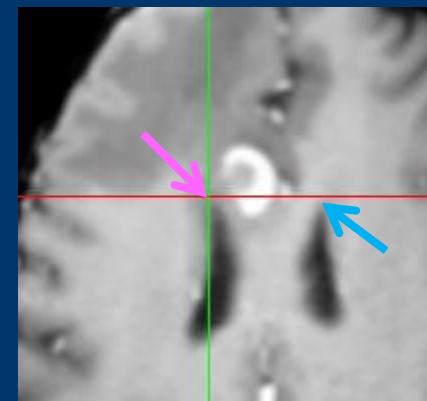
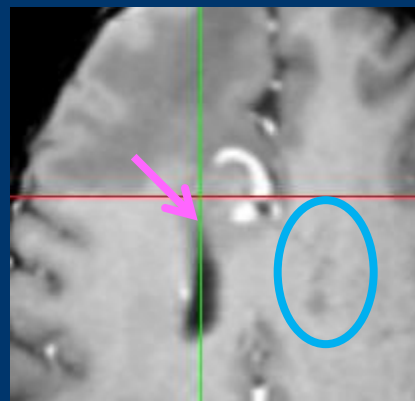
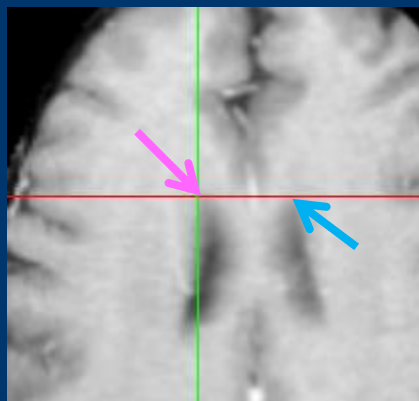
# Patient Examples: Data Preparation

- Patient 1 (4 images): Right frontal lesion, Gamma knife treatment, resection
- Patient 2 (4 images): Right parietal lesion, pre-treatment studies, resection, post-op
- Pre-processing

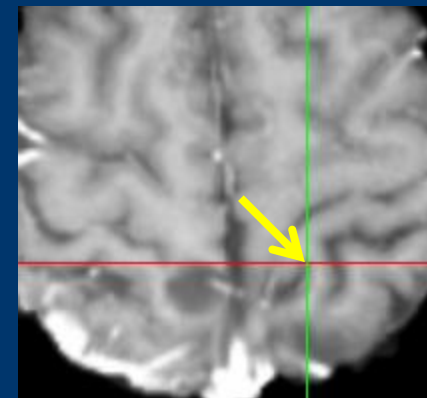
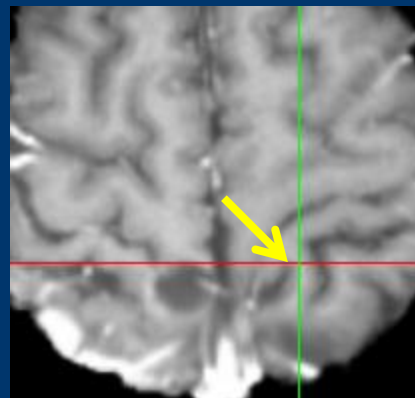
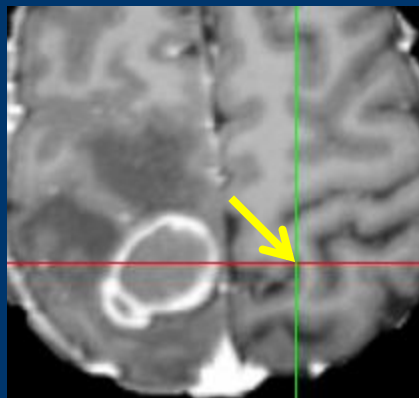


# Patient Examples: Sample Results

*Patient 1*



*Patient 2*



*Time 1*

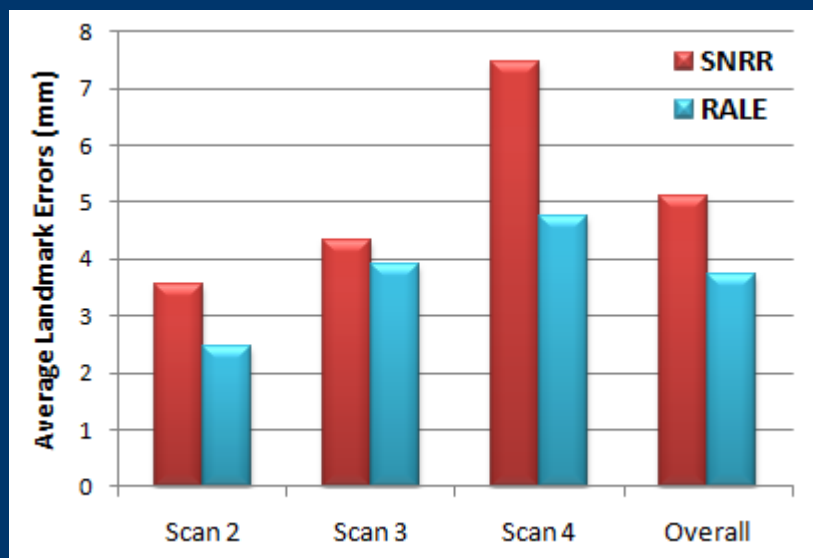
*SNRR*

*RALE*

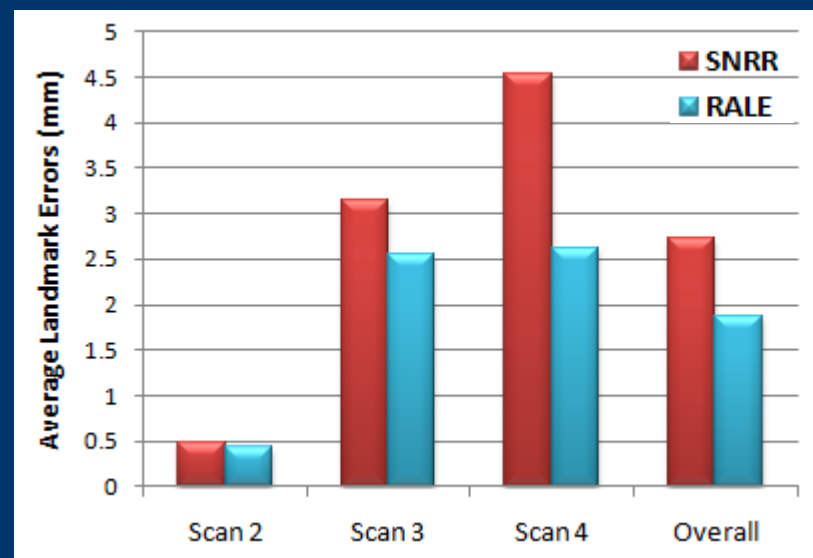
# Patient Examples: Registration Errors

Measured 7 Landmarks Highly Affected by Lesion in Each Patient

*Patient 1*



*Patient 2*



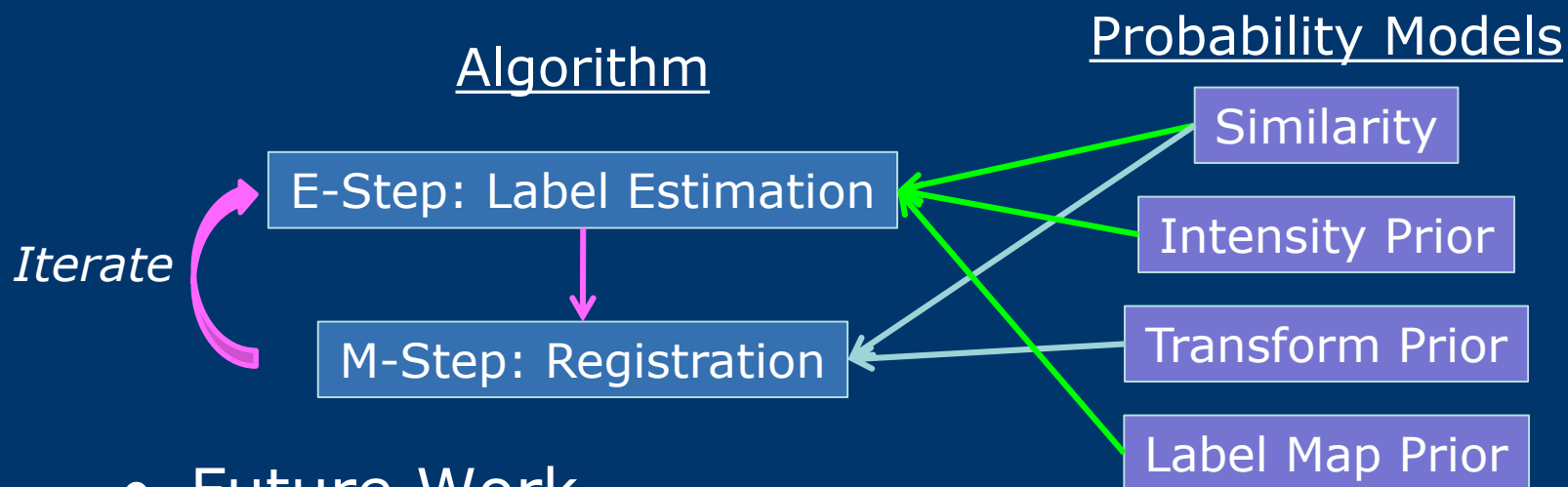
Overall Average Errors	SNRR	3.91 mm
	RALE	2.79 mm

\* RALE significantly increased accuracy ( $p < 2e-6$ )

# Conclusions and Future Work

- Contributions

- Registration method for aligning longitudinal brain tumor treatment images



- Future Work

- Difficulty distinguishing abnormal regions  
→ May include prior spatial information



# Thank You!

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- Contact: [nicha.chitphakdithai@yale.edu](mailto:nicha.chitphakdithai@yale.edu)



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