

Intraoperative Registration of Pathology for Adjuvant Postoperative Radiotherapy

Project 4. Seminar Presentation

Steven Lin

Team: Kareem Fakhoury, Matthew Hauser

Mentors: Dr. Harry Quon, Dr. Jeremy Richmon, Dr. Junghoon Lee

Overview

- Paper 1:
 - J. B. A. Maintz and M. A. Viergever. **A survey of medical image registration.** *Med. Image Anal.*, 2(1):1 –36, 1998.
- Paper 2:
 - S. Klein, M. Staring, and J. P. W. Pluim. **Evaluation of optimisation methods for nonrigid medical image registration using mutual information and B-splines.** *IEEE Trans. Image Process.*, 16(12):2879 – 2890, December 2007.



Paper 1: A survey of medical image registration

- *Purpose:*
 - A survey to compare and categorize image registration techniques.
 - Characterize registration models based on nine criteria.

Classification of Registration Methods

- I. Dimensionality
- II. Nature of Registration
- III. Nature of Transformation
- IV. Domain of Transformation
- V. Interaction
- VI. Optimization Procedure
- VII. Modalities Involved
- VIII. Subject
- IX. Object

Classification of Registration Methods

- I. Dimensionality
- II. Nature of Registration
- III. Nature of Transformation
- IV. Domain of Transformation
- V. Interaction
- VI. Optimization Procedure
- VII. Modalities Involved
- VIII. Subject
- IX. Object



Dimensionality

- Registered dimensions:
 - 2-D – 2-D
 - 2-D – 3-D
 - 3-D – 3-D
 - Can also depend on time, which would be useful for a progression study
- ***Project Relevance: 3-D to 3-D, no time series necessary.***



Classification of Registration Methods

- I. Dimensionality
- II. Nature of Registration**
- III. Nature of Transformation
- IV. Domain of Transformation
- V. Interaction
- VI. Optimization Procedure
- VII. Modalities Involved
- VIII. Subject
- IX. Object



Nature of Registration - Extrinsic

- Rely on artificial objects
- Objects may be:
 - Invasive (e.g. screw markers)
 - Non-invasive (e.g. skin markers)



Nature of Registration - Intrinsic

- Based on the image of the patient
 - *Landmark based:*
 - User identify points of visible anatomy. This means a small amount of data compare to entire image set, good for faster registration.
 - Often used in combination with other registration basis.
 - *Segmentation based:*
 - Extracted structure is fit to the second image
 - In deformable, a template is deformed to fit a second image
 - *Voxel property based:*
 - Operate by reducing image grey values, into important parameters
 - Use full image content



Nature of Registration

- **Project Relevance:**
 - *Need extrinsic markers in order to map intraoperative to pre-operative.*
 - *Dr. Lee has suggested the usage of landmark based pre-registration with other methods.*
 - *Deformable segment may involve initial pre-registration and a deformed model that is sufficiently similar to a defined template. However, this would no be an issue in our application.*
 - *Dr. Lee suggested local constraints to eliminate registration errors that would come from very similar anatomy.*
 - *Full image voxel based registration may be of use.*



Classification of Registration Methods

- I. Dimensionality
- II. Nature of Registration
- III. Nature of Transformation
- IV. Domain of Transformation
- V. Interaction
- VI. Optimization Procedure
- VII. Modalities Involved
- VIII. Subject
- IX. Object

Nature of Transformation

- Rigid – only translation/rotation
- Affine – maps parallel lines onto parallel lines
- Projective – maps lines onto lines
- Curved (elastic) – lines onto curves



Domain of Transformation

- Local transformation – composite of multiple transformation on sub-images
- Global transformation – single transformation on image
- *Project Relevance – we will most likely adopt composite of curved and rigid local transformations based on where soft tissue is removed*



Classification of Registration Methods

- I. Dimensionality
- II. Nature of Registration
- III. Nature of Transformation
- IV. Domain of Transformation
- V. Interaction**
- VI. Optimization Procedure
- VII. Modalities Involved
- VIII. Subject
- IX. Object



Interaction

- Interactive – user guided
- Semi-automatic – may be user initialized, steered or both
- Automatic

- *Project Relevance - Elastix, the package where this paper is referenced, uses a semi-automatic registration method. The algorithm is selected but the user can alter the parameters*



Classification of Registration Methods

- I. Dimensionality
- II. Nature of Registration
- III. Nature of Transformation
- IV. Domain of Transformation
- V. Interaction
- VI. Optimization Procedure**
- VII. Modalities Involved
- VIII. Subject
- IX. Object

Optimization Computed

- Parameters computed through an explicit fashion
- Parameters searched by optimizing a cost function
- *Project Relevance: Elastix uses an algorithm that searches for the parameters through maximizing a mutual information similarity measure.*



Classification of Registration Methods

- I. Dimensionality
- II. Nature of Registration
- III. Nature of Transformation
- IV. Domain of Transformation
- V. Interaction
- VI. Optimization Procedure
- VII. Modalities Involved**
- VIII. Subject
- IX. Object



Modalities

- Monomodality – registration between same type of image (e.g. CT-CT)
- Multimodality - registration between different types of image (e.g. CT-MR)
- *Project Relevance: The core of this project is in the monomodality CT-CT registration. The registration of intraoperative to pre-operative is done with Polaris-CT.*



Classification of Registration Methods

- I. Dimensionality
- II. Nature of Registration
- III. Nature of Transformation
- IV. Domain of Transformation
- V. Interaction
- VI. Optimization Procedure
- VII. Modalities Involved
- VIII. Subject**
- IX. Object**



Subject and Object

- *Project Relevance: Our project involves a intrasubject registration, where the object is the head.*

Discussion of Paper 1

- Useful in identifying the algorithms to look for
- Explain certain functionalities in Elastix
- Provide future research papers and methods



Paper 2: Evaluation of optimisation methods for nonrigid medical image registration using mutual information and B-splines

- *Purpose:*

- Introduce registration techniques that maximize mutual information and constructs a deformation field with cubic B-spline
- Compare accuracy, precision and convergence properties of eight methods

Optimization

- Family of registration operate by maximizing the mutual information matrix:

$$\hat{\mu} = \arg \min_{\mu} C(\mu; I_F, I_M)$$

- $\mu \equiv$ B-spline coefficients that defines a deformation field
- $C \equiv$ mutual information similarity metric
- I_F is the fixed image
- I_M is the moving, deformed image



Mutual Information

- Measures information that two random variables share
- Definition of mutual information for discrete random variables

$$MI(\boldsymbol{\mu}; I_F, I_M) = \sum_{m \in L_M} \sum_{f \in L_F} p(f, m; \boldsymbol{\mu}) \times \log_2 \left(\frac{p(f, m; \boldsymbol{\mu})}{p_F(f)p_M(m; \boldsymbol{\mu})} \right)$$



Optimization Continued

- μ can be found using an iterative optimization

$$\boldsymbol{\mu}_{k+1} = \boldsymbol{\mu}_k + a_k \mathbf{d}_k, \quad k = 0, 1, 2, \dots$$

- a_k is a gain factor, that controls the step size
- d_k is the search direction

- Algorithms vary based on the computation of a_k and d_k
 - Deterministic gradient-based
 - Stochastic gradient-based
 - Evolution strategy



Gradient Descent (GDD and GDL)

$$\boldsymbol{\mu}_{k+1} = \boldsymbol{\mu}_k + a_k \mathbf{d}_k, \quad k = 0, 1, 2, \dots$$

- $d_k = g(\boldsymbol{\mu}_k)$, derivative of the cost function

$$\boldsymbol{\mu}_{k+1} = \boldsymbol{\mu}_k - a_k \mathbf{g}(\boldsymbol{\mu}_k)$$

- GDD: $a_k = \frac{a}{(k+A)^\alpha}$, with user a , A , α
- GDL: a_k determined through Moré-Thuente routine



Quasi-Newton

- Based on Newton-Raphson algorithm

$$\boldsymbol{\mu}_{k+1} = \boldsymbol{\mu}_k - a_k L_k \mathbf{g}(\boldsymbol{\mu}_k).$$

- $L_k \approx [H(\boldsymbol{\mu}_k)]^{-1}$, approximation of the inverse Hessian of the cost function. Can be found using the LBFGS method
- a_k is found using Moré-Thuente routine
 - Inexact line search routine that finds a_k so that it satisfies the strong Wolfe conditions

$$\begin{aligned} C(\boldsymbol{\mu}_{k+1}) &\leq C(\boldsymbol{\mu}_k) + c_1 a_k \mathbf{d}_k^T \mathbf{g}(\boldsymbol{\mu}_k) \\ \left| \mathbf{d}_k^T \mathbf{g}(\boldsymbol{\mu}_{k+1}) \right| &\leq c_2 \left| \mathbf{d}_k^T \mathbf{g}(\boldsymbol{\mu}_k) \right| \end{aligned}$$

Nonlinear Conjugate Gradient

- Direction determined through linear combination of the gradient and the previous direction

$$\mathbf{d}_k = -\mathbf{g}(\boldsymbol{\mu}_k) + \beta_k \mathbf{d}_{k-1}.$$

- The factor β_k is computed through

$$\beta_k = \max(0, \min(\beta_k^{\text{HS}}, \beta_k^{\text{DY}}))$$

$$\text{Dai - Yuan : } \beta_k^{\text{DY}} = \frac{\mathbf{g}_k^T \mathbf{g}_k}{\mathbf{d}_{k-1}^T (\mathbf{g}_k - \mathbf{g}_{k-1})}$$

$$\text{Hestenes - Stiefel : } \beta_k^{\text{HS}} = \frac{\mathbf{g}_k^T (\mathbf{g}_k - \mathbf{g}_{k-1})}{\mathbf{d}_{k-1}^T (\mathbf{g}_k - \mathbf{g}_{k-1})}$$



Stochastic Gradient Descent (KW, SP, RM)

$$\boldsymbol{\mu}_{k+1} = \boldsymbol{\mu}_k - \alpha_k \tilde{\mathbf{g}}_k$$

- $\tilde{\mathbf{g}}_k$ is an approximation of the gradient
- *Kiefer-Wolfowitz*: Finite difference approximation

$$[\tilde{\mathbf{g}}_k]_i = \frac{\mathcal{C}(\boldsymbol{\mu}_k + c_k \mathbf{e}_i) - \mathcal{C}(\boldsymbol{\mu}_k - c_k \mathbf{e}_i)}{2c_k}$$

$$[\tilde{\mathbf{g}}_k]_i = \frac{\tilde{\mathcal{C}}_{ki}^+ - \tilde{\mathcal{C}}_{ki}^-}{2c_k}$$

$$\tilde{\mathcal{C}}_{ki}^+ = \mathcal{C}(\boldsymbol{\mu}_k + c_k \mathbf{e}_i) + \varepsilon_{ki}^+$$

$$\tilde{\mathcal{C}}_{ki}^- = \mathcal{C}(\boldsymbol{\mu}_k - c_k \mathbf{e}_i) + \varepsilon_{ki}^-$$

- *Simultaneous Perturbation:*

$$[\tilde{\mathbf{g}}_k]_i = \frac{\tilde{\mathcal{C}}_k^+ - \tilde{\mathcal{C}}_k^-}{2c_k[\Delta_k]_i}$$

$$\tilde{\mathcal{C}}_k^+ = \mathcal{C}(\boldsymbol{\mu}_k + c_k \boldsymbol{\Delta}_k) + \varepsilon_k^+$$

$$\tilde{\mathcal{C}}_k^- = \mathcal{C}(\boldsymbol{\mu}_k - c_k \boldsymbol{\Delta}_k) + \varepsilon_k^-$$

- *Robbins-Monro:* Assume the approximation of the cost function

$$\tilde{\mathbf{g}}_k = \mathbf{g}(\boldsymbol{\mu}_k) + \boldsymbol{\varepsilon}_k$$

- Approximation done with randomly selected subset of voxels at each iteration

Evolution Strategy

- Based on natural selection
- Each iteration has three phases: *Offspring generation*, *selection*, *recombination*
- *Offspring generation*: set of λ direction is found in $\mathcal{N}(0, C_k)$.
- *Selection*: the P directions that has the lowest value in the cost function, $\mathcal{C}(\mu_k + a_k \mathbf{d}_k^{(\ell)})$, are found
- *Recombination*: All P directions are summed with weighing factors

$$\mathbf{d}_k = \sum_{p=1}^P w_p \mathbf{d}_k^{(p;\lambda)}$$



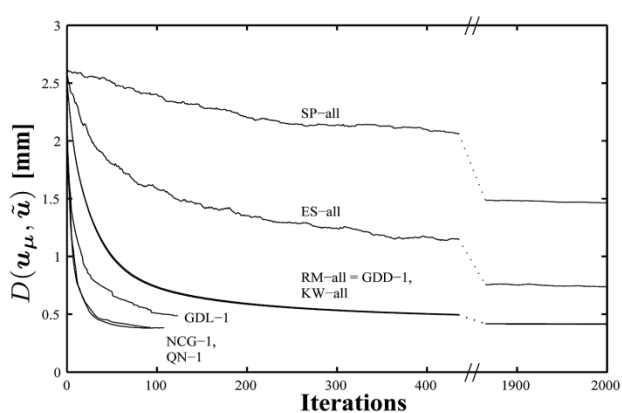
Experiments and Results

- Four CT of the heart
- Known deformation field \tilde{u} generated with randomly placed Gaussian blobs.
- Accuracy is evaluated using the following:

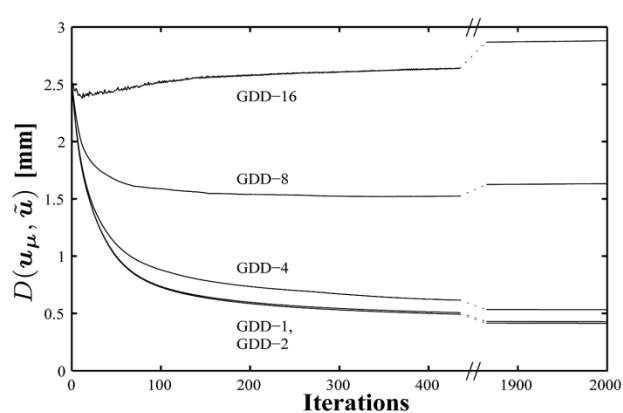
$$D(\mathbf{u}_1, \mathbf{u}_2) = \frac{1}{|I_F|} \sum_{\mathbf{x}_i \in I_F} \|\mathbf{u}_1(\mathbf{x}_i) - \mathbf{u}_2(\mathbf{x}_i)\|$$

- Overall, RM is the most accurate.

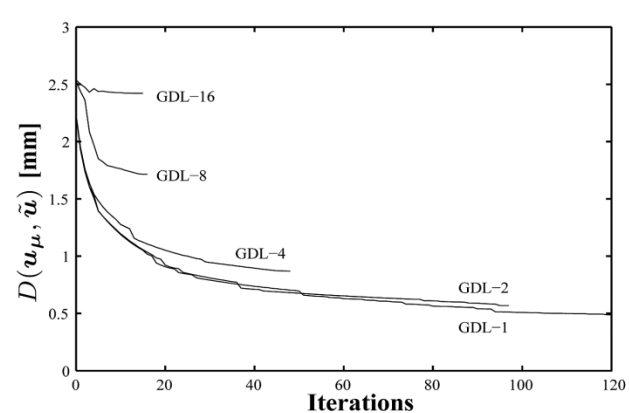




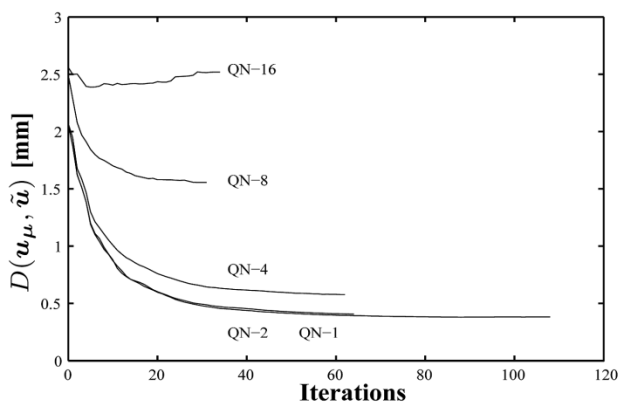
(a) All methods without subsampling



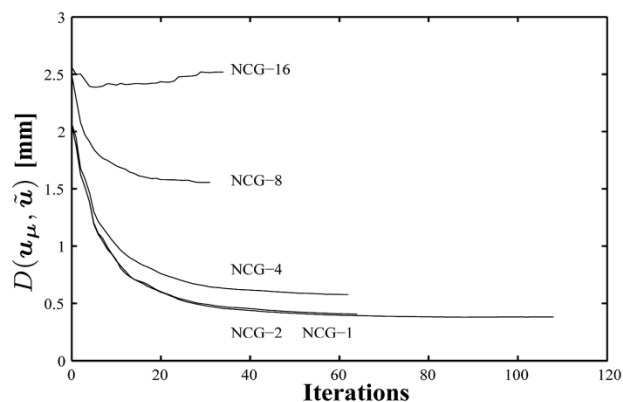
(b) GDD



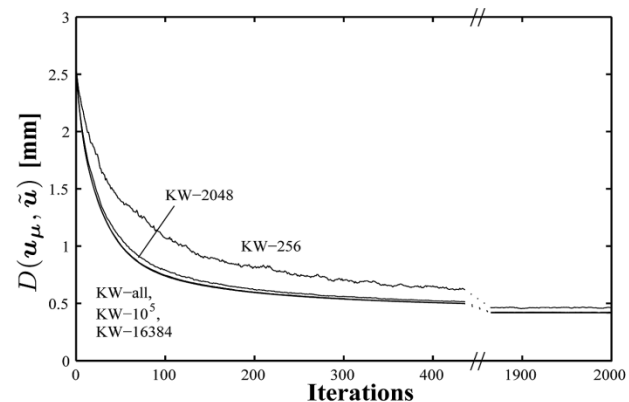
(c) GDL



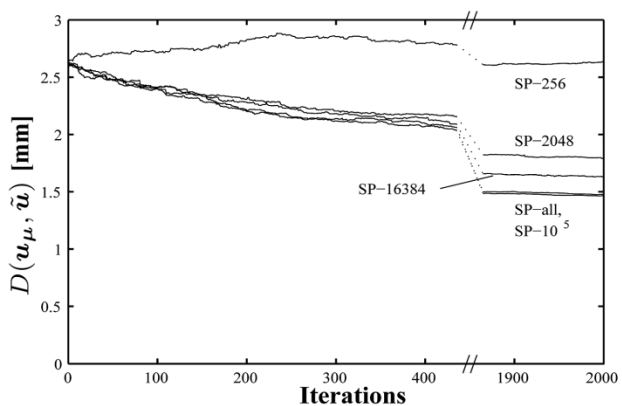
(d) QN



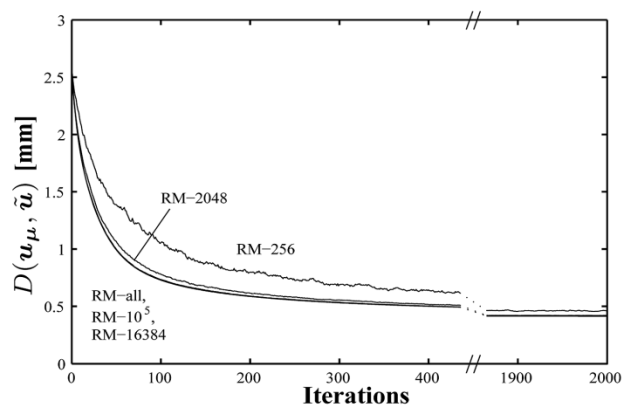
(e) NCG



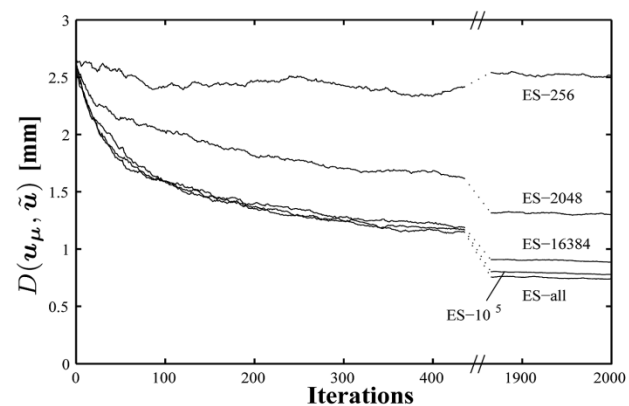
(f) KW



(g) SP



(h) RM



(i) ES

Experiments and Results

- Five patients, chest CT's were taken months apart.
- Images were registered.
- Precision was measured with

$$D(\mathbf{u}_1, \mathbf{u}_2) = \frac{1}{|I_F|} \sum_{\mathbf{x}_i \in I_F} \|\mathbf{u}_1(\mathbf{x}_i) - \mathbf{u}_2(\mathbf{x}_i)\|$$

- Dice similarity index is used to measure accuracy.

$$\text{overlap} = \frac{2 |V_1 \cap V_2|}{|V_1| + |V_2|} \cdot 100\%$$

- RM was found to be the best method.



	CT follow-up chest ($t_g \approx 220$ s.)				MR BFFE prostate ($t_g \approx 56$ s.)						MR T1-T2 prostate ($t_g \approx 9$ s.)		
	time	overlap	precision	effect \mathcal{R}	time	overlap	precision	effect \mathcal{R}	overlap*	precision*	time	precision	effect \mathcal{R}
	avg [t_g]	avg \pm sd [%]	avg \pm sd [mm]	avg \pm sd [mm]	avg [t_g]	avg \pm sd [%]	avg \pm sd [mm]	avg \pm sd [mm]	avg \pm sd [%]	avg \pm sd [mm]	avg [t_g]	avg \pm sd [mm]	avg \pm sd [mm]
rigid		36 \pm 15	9.2 \pm 7.1			37 \pm 11	3.2 \pm 1.0					2.9 \pm 0.8	
GDL-1	700	76 \pm 7	0.1 \pm 0.1	1.0 \pm 0.4	700	58 \pm 5	0.1 \pm 0.1	2.2 \pm 0.9	58 \pm 6	0.2 \pm 0.1	100	0.6 \pm 0.4	1.4 \pm 0.5
GDL-2	100	75 \pm 6	0.4 \pm 0.1	0.9 \pm 0.3	300	58 \pm 5	0.2 \pm 0.1	2.0 \pm 0.9	58 \pm 6	0.3 \pm 0.2	50	0.7 \pm 0.4	1.4 \pm 0.5
GDL-4	10	75 \pm 7	0.7 \pm 0.2	0.6 \pm 0.3	40	57 \pm 6	0.6 \pm 0.2	1.8 \pm 0.7	57 \pm 6	0.7 \pm 0.3	10	1.1 \pm 0.6	1.7 \pm 0.4
GDL-8	1	71 \pm 7	1.3 \pm 0.3	0.6 \pm 0.3	9	56 \pm 6	1.6 \pm 0.7	1.6 \pm 0.6	55 \pm 6	1.8 \pm 0.7	2	1.7 \pm 0.5	1.2 \pm 0.3
GDL-16	0.09	60 \pm 12	3.3 \pm 2.6	0.9 \pm 0.3	1	45 \pm 6	3.3 \pm 0.9	2.0 \pm 0.5	43 \pm 6	2.9 \pm 1.0	1	3.0 \pm 0.8	1.6 \pm 0.7
QN-1	200	77 \pm 7	0.0 \pm 0.0	1.7 \pm 0.8	100	58 \pm 5	0.0 \pm 0.0	4.0 \pm 2.0	58 \pm 6	0.0 \pm 0.0	60	0.0 \pm 0.0	4.4 \pm 2.1
QN-2	40	76 \pm 7	0.2 \pm 0.1	1.4 \pm 0.6	40	58 \pm 5	0.1 \pm 0.0	3.9 \pm 2.0	58 \pm 6	0.1 \pm 0.0	20	0.4 \pm 0.1	4.8 \pm 2.2
QN-4	5	75 \pm 7	0.7 \pm 0.2	1.3 \pm 0.5	8	57 \pm 5	0.6 \pm 0.6	3.4 \pm 1.7	57 \pm 6	0.5 \pm 0.2	7	1.0 \pm 0.6	4.9 \pm 1.8
QN-8	0.5	71 \pm 7	1.3 \pm 0.3	1.4 \pm 0.5	1	56 \pm 6	1.4 \pm 0.6	3.4 \pm 1.2	55 \pm 6	1.9 \pm 0.8	2	1.9 \pm 0.9	4.7 \pm 1.2
QN-16	0.1	57 \pm 7	2.8 \pm 0.9	2.4 \pm 0.8	0.2	43 \pm 5	3.5 \pm 0.9	4.0 \pm 1.2	40 \pm 8	3.2 \pm 0.7	0.3	3.9 \pm 1.0	4.3 \pm 1.6
NCG-1	300	77 \pm 7	0.1 \pm 0.0	1.4 \pm 0.6	200	58 \pm 5	0.0 \pm 0.0	3.0 \pm 1.6	58 \pm 6	0.0 \pm 0.0	70	0.2 \pm 0.2	2.8 \pm 1.5
NCG-2	40	76 \pm 7	0.2 \pm 0.1	1.3 \pm 0.6	70	58 \pm 5	0.1 \pm 0.1	2.8 \pm 1.3	58 \pm 6	0.2 \pm 0.1	30	0.5 \pm 0.3	2.8 \pm 1.4
NCG-4	5	75 \pm 7	0.7 \pm 0.2	1.2 \pm 0.6	10	57 \pm 6	0.7 \pm 0.5	2.5 \pm 1.1	57 \pm 6	0.6 \pm 0.3	7	1.1 \pm 0.6	3.4 \pm 1.2
NCG-8	0.5	71 \pm 8	1.4 \pm 0.5	1.6 \pm 0.6	2	56 \pm 5	1.5 \pm 0.6	2.3 \pm 1.0	55 \pm 6	1.7 \pm 0.7	2	1.7 \pm 0.7	2.7 \pm 0.8
NCG-16	0.07	57 \pm 9	3.4 \pm 2.3	2.8 \pm 2.3	0.5	46 \pm 6	3.3 \pm 0.9	3.0 \pm 0.8	41 \pm 10	3.3 \pm 1.0	0.2	3.6 \pm 0.9	3.2 \pm 1.1
RM-all	1000	76 \pm 7	0.2 \pm 0.1	0.6 \pm 0.3	2000	57 \pm 6	0.4 \pm 0.2	1.0 \pm 0.4	58 \pm 6	0.3 \pm 0.2	3000	0.6 \pm 0.4	0.9 \pm 0.6
RM-10 ⁵	30	76 \pm 7	0.2 \pm 0.1	0.6 \pm 0.2	200	57 \pm 6	0.4 \pm 0.2	1.0 \pm 0.4	58 \pm 6	0.3 \pm 0.2	700	0.7 \pm 0.6	0.9 \pm 0.5
RM-16384	5	76 \pm 7	0.2 \pm 0.1	0.6 \pm 0.3	30	57 \pm 6	0.4 \pm 0.2	1.0 \pm 0.4	58 \pm 6	0.3 \pm 0.2	200	0.7 \pm 0.5	0.9 \pm 0.5
RM-2048	0.6	75 \pm 7	0.5 \pm 0.1	0.8 \pm 0.3	4	57 \pm 6	0.4 \pm 0.2	1.0 \pm 0.4	58 \pm 6	0.4 \pm 0.2	30	0.7 \pm 0.5	1.0 \pm 0.5
RM-256	0.08	58 \pm 5	2.6 \pm 0.6	5.6 \pm 1.1	0.5	57 \pm 6	0.7 \pm 0.4	1.1 \pm 0.4	54 \pm 8	1.3 \pm 0.8	4	1.6 \pm 0.8	2.0 \pm 1.0

Discussion of Paper 2

- Paper has demonstrated that the Robbins-Monro method may be the most valuable method.
- Quasi-Newton and Nonlinear Conjugate Gradient method may be used, with lower subsampling factors.
- Useful paper, in both understanding Elastix and future research.



Citations

- J. B. A. Maintz and M. A. Viergever. A survey of medical image registration. *Med. Image Anal.*, 2(1):1 –36, 1998.
- S. Klein, M. Staring, and J. P. W. Pluim. Evaluation of optimisation methods for nonrigid medical image registration using mutual information and B-splines. *IEEE Trans. Image Process.*, 16(12):2879 – 2890, December 2007.

