

# Thesis Proposal

## Coupled Dictionary Learning for Image Analysis

Tian Cao

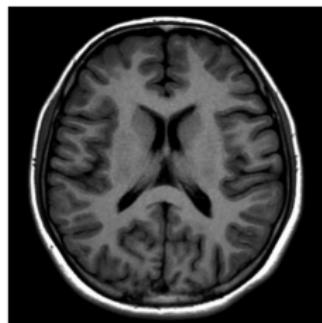
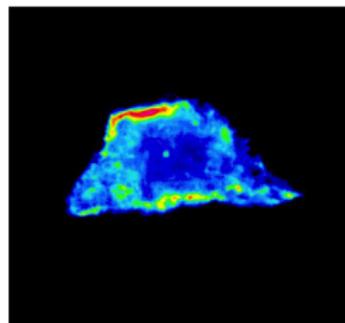
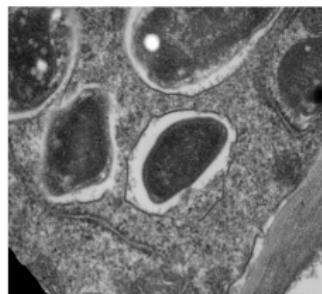
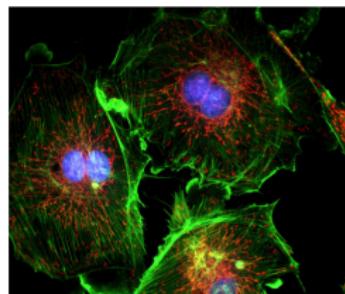
Department of Computer Science  
University of North Carolina at Chapel Hill

## Motivations

- Modern imaging technologies allow us to visualize various objects

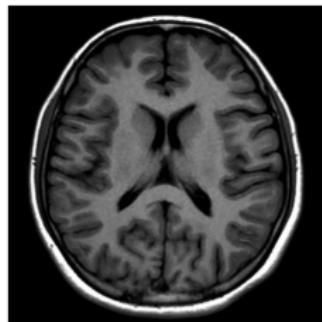
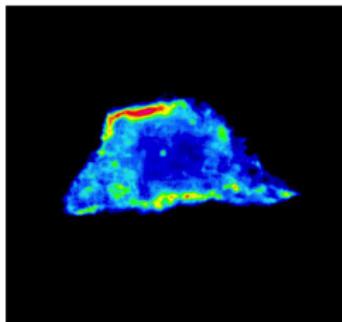
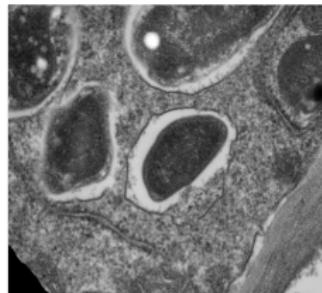
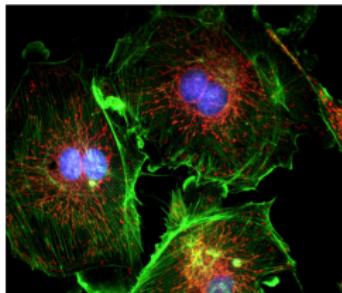
# Motivations

- Modern imaging technologies allow us to visualize various objects



# Motivations

- Modern imaging technologies allow us to visualize various objects
- Common problem in image analysis is how to relate the information from different modalities/sources



# Multi-modal Image Registration

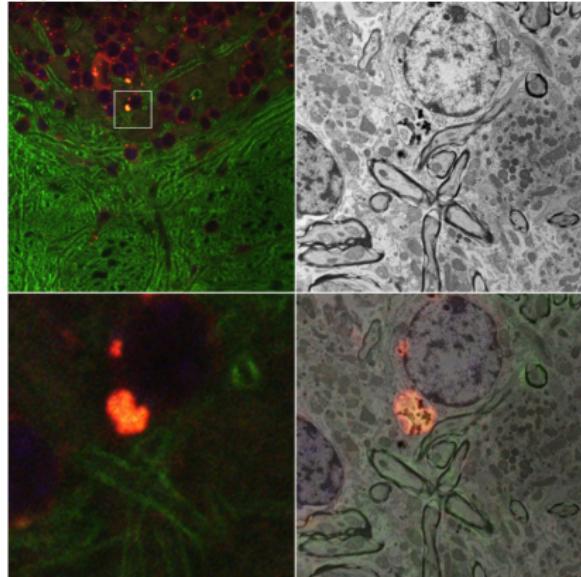
## Correlative Microscopy

- combination of different microscopy technologies
  - Confocal
  - Transmission Electron Microscopy (TEM)

Requires image registration for joint analysis

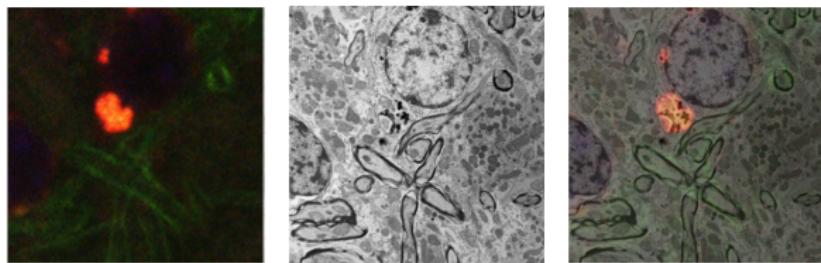
Relating image appearances from different modalities

## Confocal      TEM



# Image Registration

Image Registration is the process of estimating a spatial transformation between two images



(a) Source image

(b) Target image

(c) Registration result

Figure: Image Registration Example

# Deformation Estimation

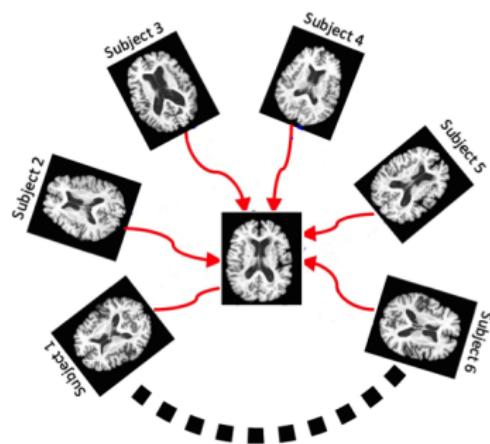
Estimating deformations to register subject images to a common reference image (atlas)

- Brain Images Analysis

Learning based approach

- estimating deformations from image appearances

Relating deformations with image appearances



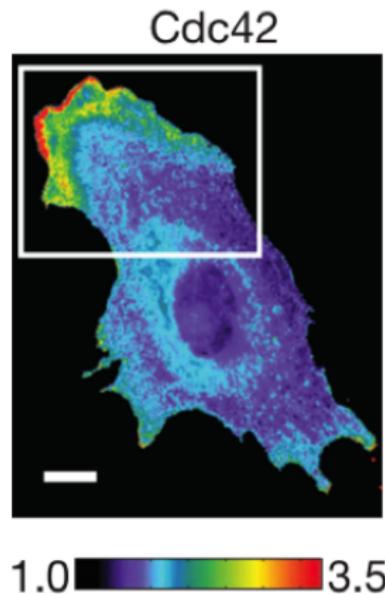
# Coordination between GTPase Activations and Cell Movements in Mouse Embryonic Fibroblasts

GTPases are a family of enzymes that can bind Guanosine triphosphate(GTP)

Establishing correspondences between GTPase activations and cell movements

- crucial for understand cell dynamics

Relating spatiotemporal information



## Similarities between The Three Examples

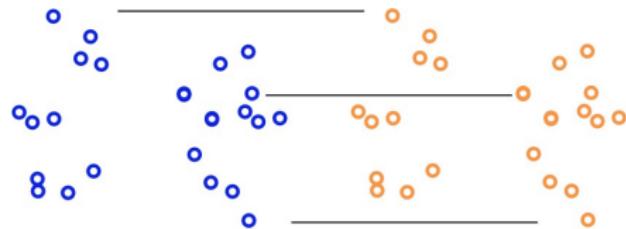
Examples:

- multi-modal registration for correlative microscopy
- deformation estimation from image appearances
- coordination between GTPase activations and cell movements

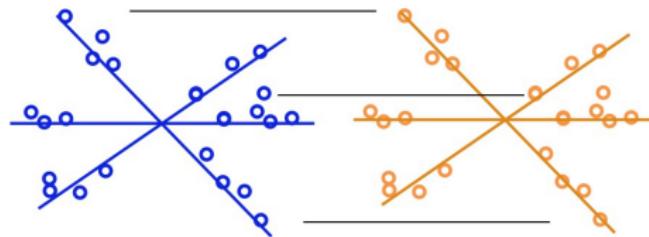
Similarities:

- relating data from two different spaces
  - spatial data: appearances
  - spatial data: appearances and deformations
  - spatiotemporal data: GTP activations and cell movements
- can be solved by modeling the relationship between the data in two spaces

# My Solution

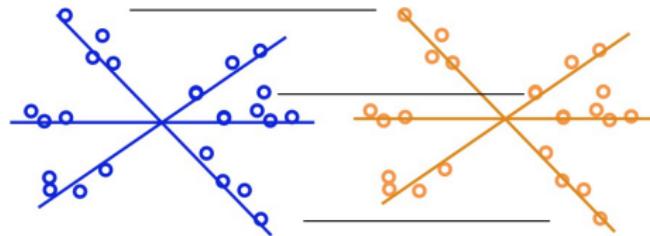


## My Solution



- Learning a joint basis for compact representation of two spaces

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- Learning a joint basis for compact representation of two spaces
- Joint basis is learned by coupled dictionary learning

# Thesis

- Learning a coupled basis for the compact representation of two spaces can be achieved by **coupled dictionary learning** (CDL).

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- Such dictionaries can be learned to capture **appearance differences** of different imaging modalities, dependencies between **image appearances and deformations** as well as the **spatiotemporal patterns** for cell signaling and boundary protrusions and retractions.

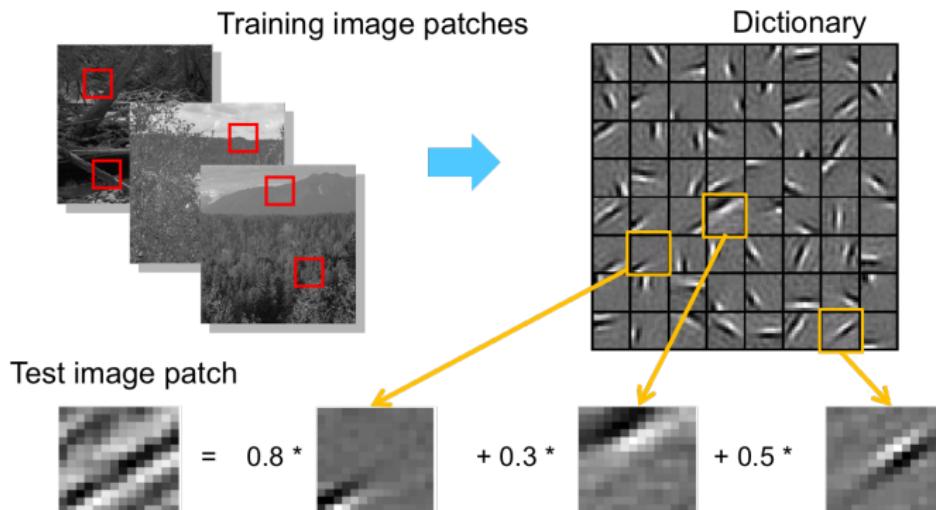
# Thesis

- Learning a coupled basis for the compact representation of two spaces can be achieved by **coupled dictionary learning** (CDL).
- Such dictionaries can be learned to capture **appearance differences** of different imaging modalities, dependencies between **image appearances and deformations** as well as the **spatiotemporal patterns** for cell signaling and boundary protrusions and retractions.
- To account for data inconsistencies, a robust coupled dictionary can be obtained based on a **probabilistic dictionary model**.

# Outline

- Background
- CDL for Multi-modal Image registration
- CDL for Deformation Estimation
- CDL for GTPase Activations and Cell Protrusions
- Robust CDL
- Summary and Future Work

# Sparse Representation Models (SRM)



- Dictionary is a basis to represent the data in a space
- Each coefficient have few non-zero values (sparsity)

## Sparse Representation Model (SRM)

Suppose  $D$  is pre-defined, sparse representation model includes solving the optimization problem

$$\hat{\alpha} = \arg \min_{\alpha} \frac{1}{2} \|x - D\alpha\|_2^2 + \lambda \|\alpha\|_p$$

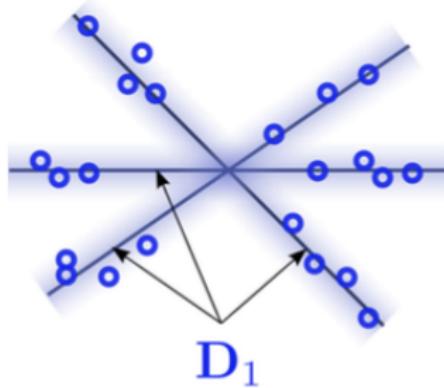
Usual sparsity-inducing regularizers

- $p = 0$ :  $\ell_0$  norm, number of non-zero elements, non-convex, greedy algorithm
- $p = 1$ :  $\ell_1$  norm, relaxation of  $\ell_0$  norm, convex optimization (lasso)

# Dictionary Learning

$$(\hat{\alpha}, \hat{D}) = \arg \min_{\alpha, D} \sum_{i=1}^m \frac{1}{2} \|x_i - D\alpha_i\|_2^2 + \lambda \|\alpha_i\|_p$$

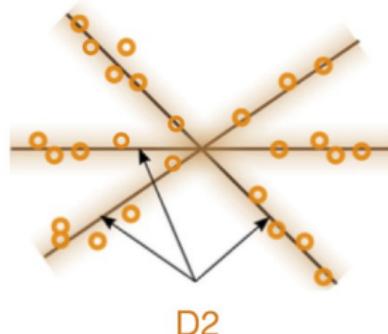
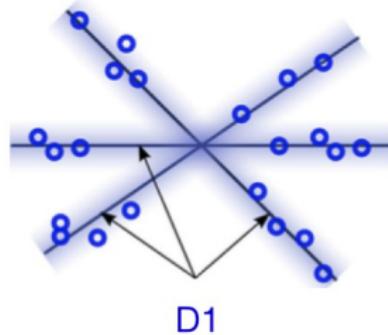
- Atoms satisfy  $\|d_i\|_2 \leq 1$
- Usual Sparsity-inducing regularizes
  - $\ell_0$  norm:  $\|\alpha_i\|_0$
  - $\ell_1$  norm:  $\|\alpha_i\|_1$



# Coupled Dictionary Learning

$$(\hat{\alpha}, \hat{D}^*) = \arg \min_{\alpha, D^*} \sum_{i=1}^m \frac{1}{2} \|x_i^* - D^* a_i\|_2^2 + \lambda \|a_i\|_1$$

- $x_i^* = [x_i^{1T}, x_i^{2T}]^T$ ,
- $D_i^* = [D_i^{1T}, D_i^{2T}]^T$
- Atoms satisfy  $\|d_i^*\|_2 \leq 1$
- Use a single  $\alpha$  to enforce the correspondences between data in two spaces



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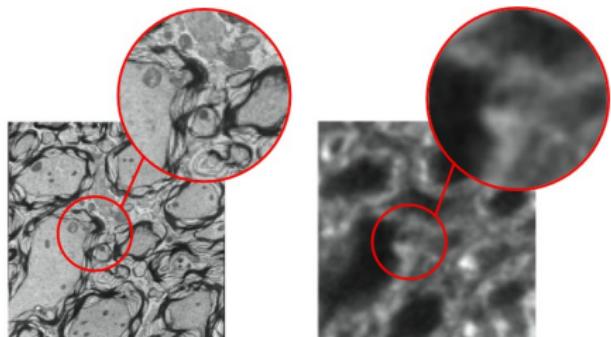
# Multi-modal Image Registration

## Challenges

- different appearances between different image modalities

## Previous Methods

- distance measure: mutual information
- change image appearance [S. Yang 2011]
- convert image modality [W. Wein 2008]
- MR image synthesis [S. Roy 2011]



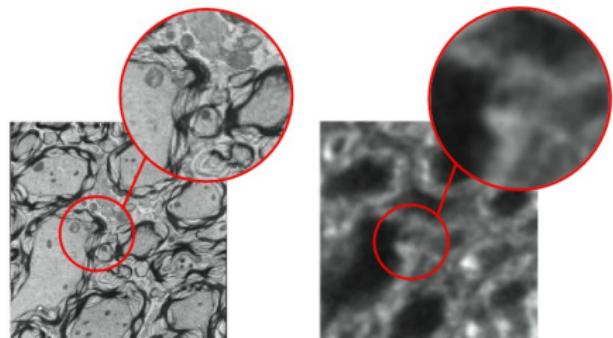
# Multi-modal Image Registration

## Challenges

- different appearances between different image modalities

## My method

- image analogies (IA)
- simplify a multi-modal registration problem to a mono-modal one



# Image Analogies Problem

## Training images

- image pairs  $A$  and  $A'$
- $A'$  is a “filtered” version of  $A$

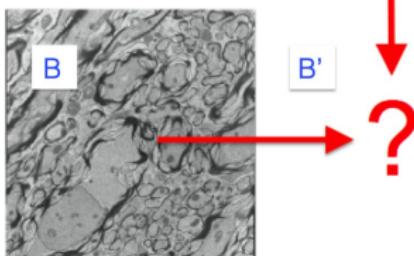
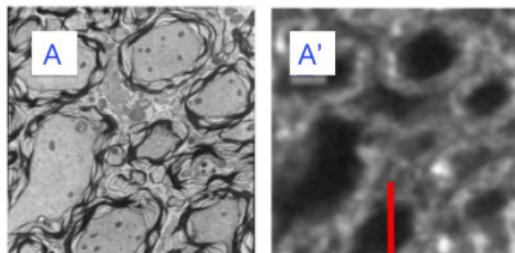
## Input image

- source image  $B$

## Output image

- predicted “filtered” image  $B'$ , based on the relationship between  $A$  and  $A'$

## Training Images

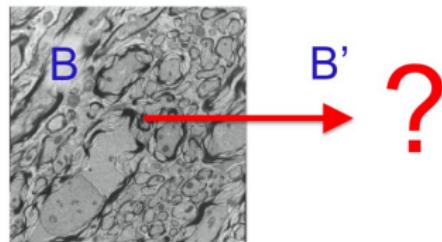
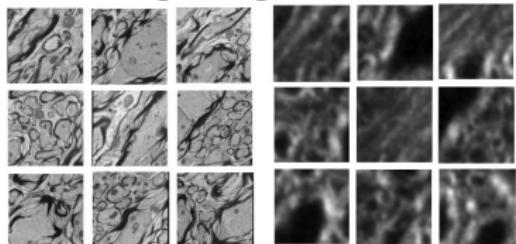


Input Image      Predicted Image

# Standard Image Analogies (IA)

- Generate image patch pairs from  $A$  and  $A'$

Training Image Patches



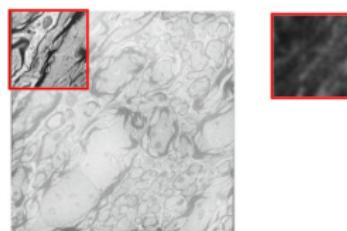
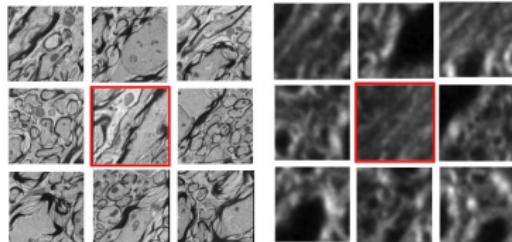
Input Image

Predicted Image

# Standard Image Analogies (IA)

- Generate image patch pairs from  $A$  and  $A'$
- Reconstruct source image  $B$  by searching the closest patches in training set
- Predict  $B'$  based on the found patches

Training Image Patches



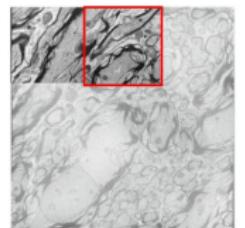
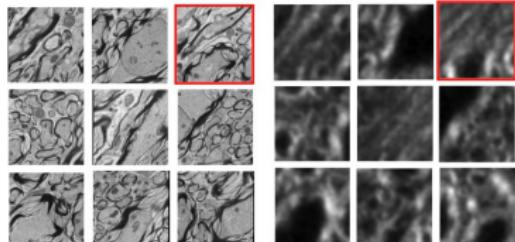
Input Image

Predicted Image

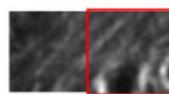
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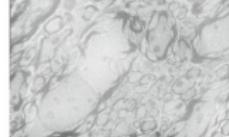
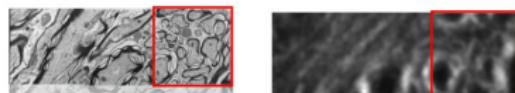
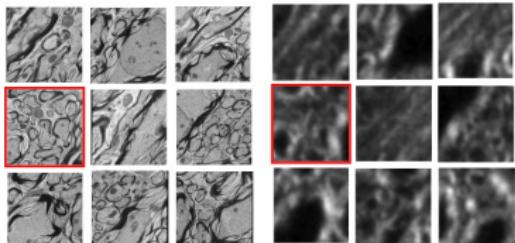


Predicted Image

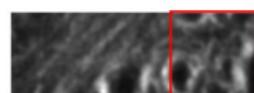
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Training Image Patches



Input Image

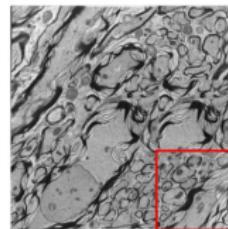
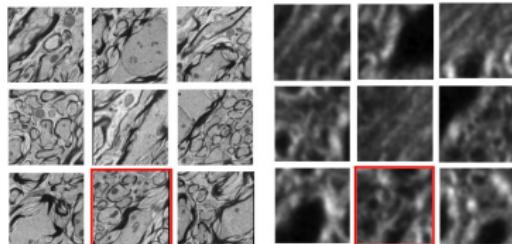


Predicted Image

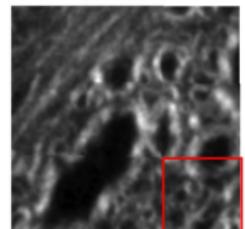
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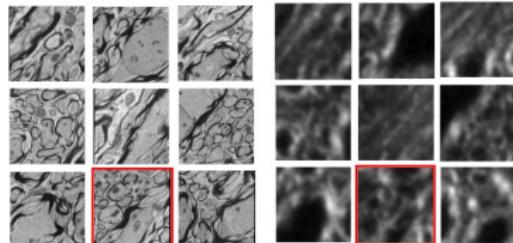
Input Image



Predicted Image

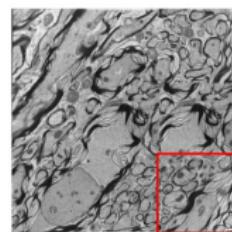
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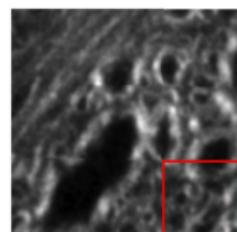


Limitation:

- nearest neighbor search in a large training set is computationally expensive



Input Image



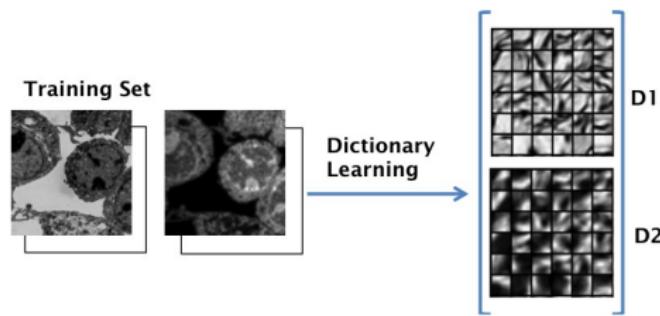
Predicted Image

## Proposed Method

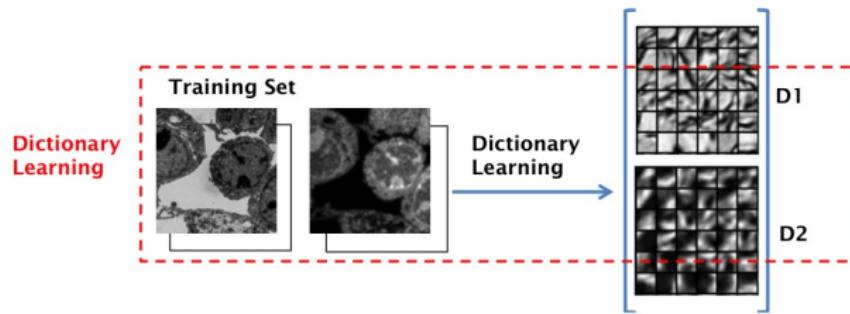
A general framework for image analogies based on

- coupled dictionary learning for **image appearances from different modalities**
- sparse representation model

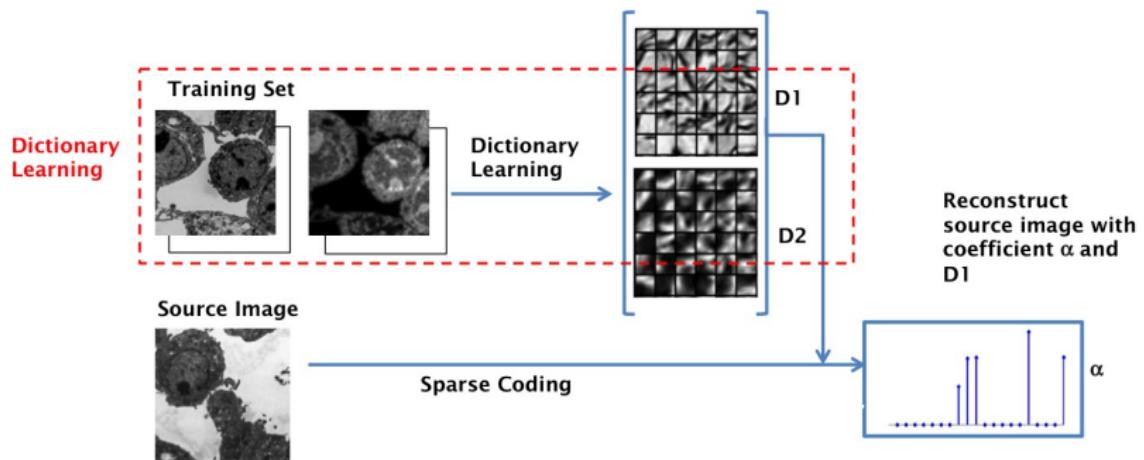
## Framework of Proposed Method



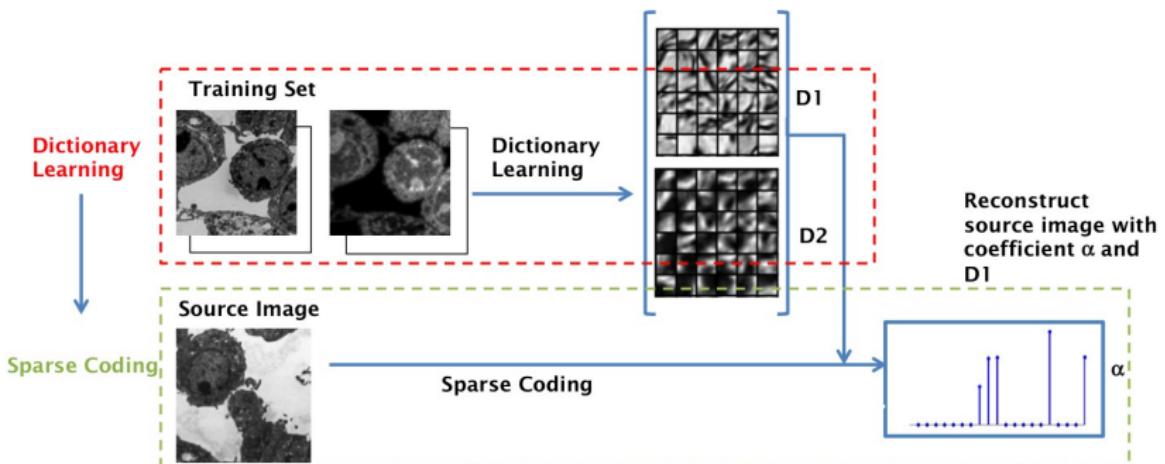
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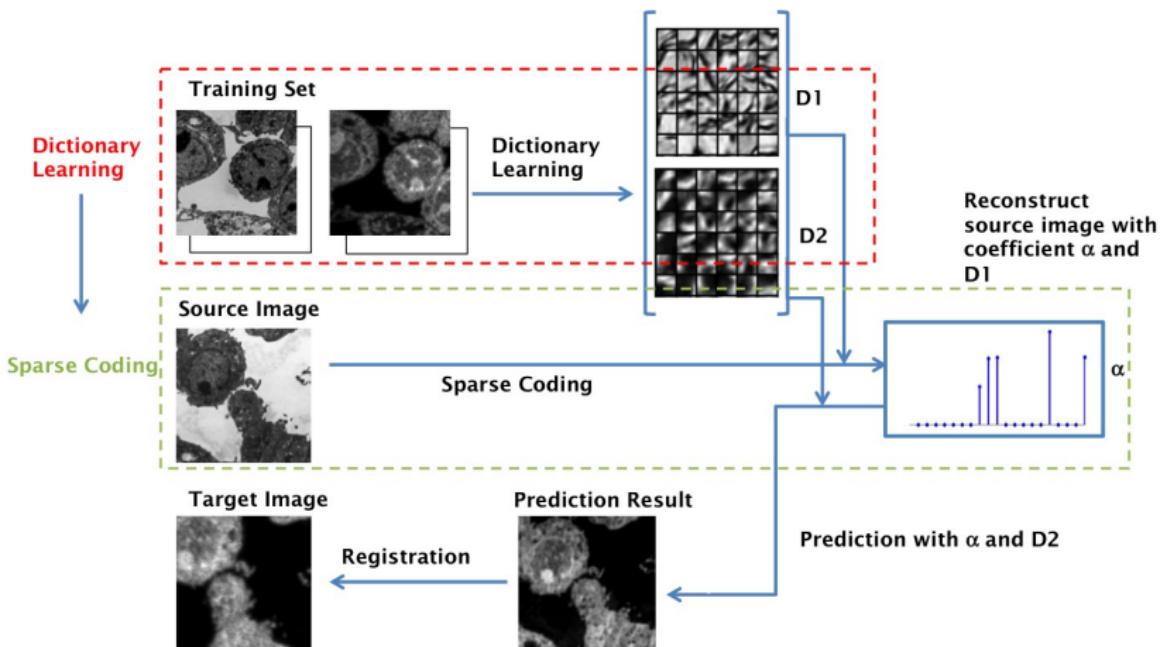
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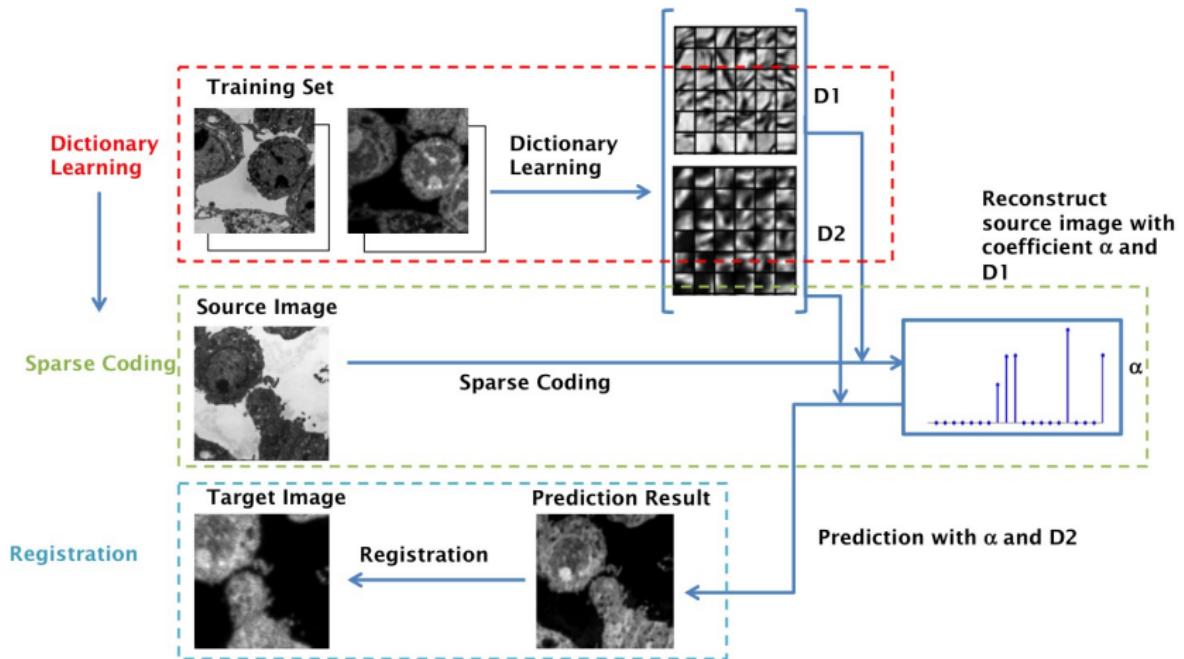
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# Framework of Proposed Method



## Formulation of Image Analogies Problem with SRM

- $f_1$  is the input image  $B$ ,  $f_2$  is the corresponding image  $B'$
- $(D_1, D_2)$  are learned coupled dictionary
- $u_1, u_2$  are reconstructed images of  $f_1, f_2$  respectively

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- $R_i u$  selects  $i$ th image patch in  $u$

$$E(u_1, u_2, \{\alpha_i\}) =$$

$$\frac{\gamma}{2} \|u_1 - f_1\|_2^2 + \frac{1}{N} \left( \frac{1}{2} \sum_{i=1}^N \|R_i \begin{pmatrix} u_1 \\ u_2 \end{pmatrix} - \begin{pmatrix} D_1 \\ D_2 \end{pmatrix} \alpha_i\|_2^2 + \lambda \|\alpha_i\|_1 \right)$$

• Similarity

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- Similarity
- Reconstruction error

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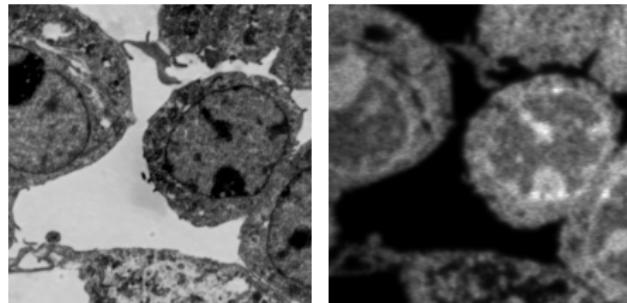
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- Similarity
- Reconstruction error
- Regularization

## Results

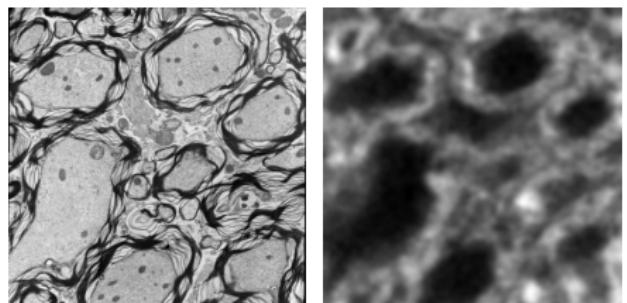
Eight pairs of 2D correlative Scanning Electron Microscopy (SEM)/Confocal images

- with gold bead fiducials
- use landmark based registration for validation

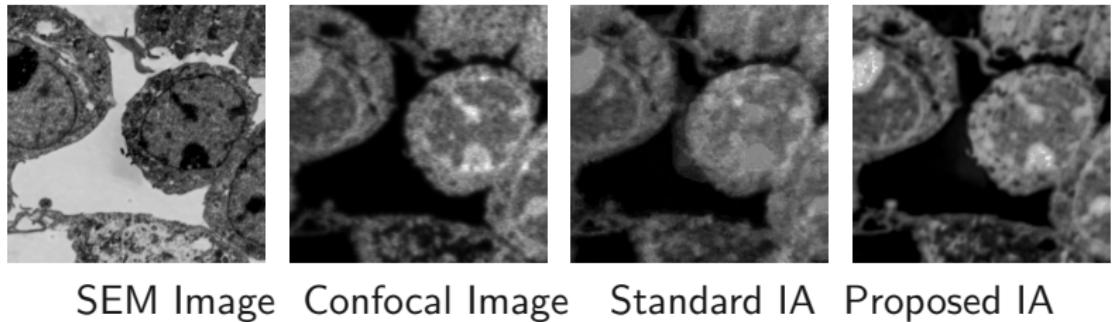


Six pairs of 2D correlative TEM/Confocal images

- without fiducials
- manually choose landmarks for validation



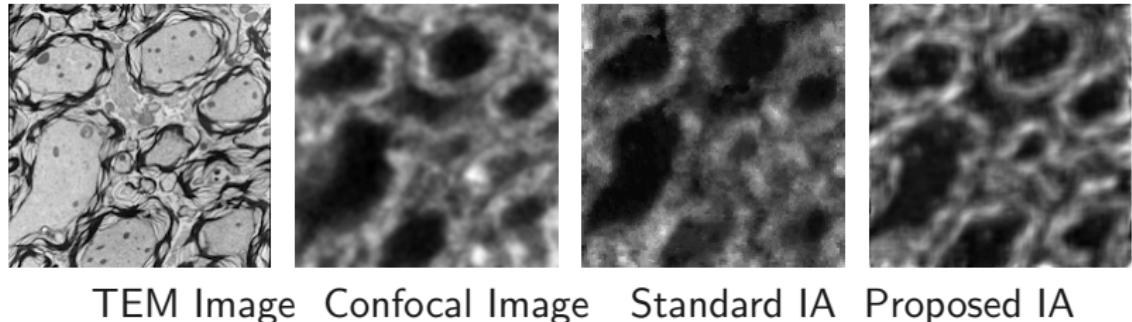
## Image Analogies Results



SEM Image Confocal Image Standard IA Proposed IA

Method	mean	std	p-value
Proposed IA	$1.52 \times 10^5$	$5.79 \times 10^4$	<b>0.0002</b>
Standard IA	$2.83 \times 10^5$	$7.11 \times 10^4$	

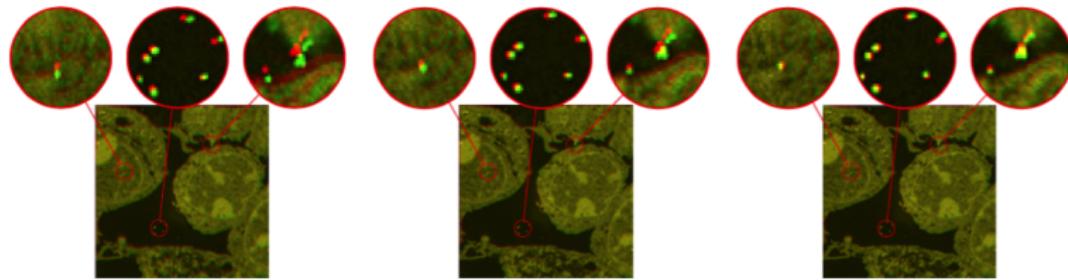
## Image Analogies Results



TEM Image Confocal Image Standard IA Proposed IA

Method	mean	std	p-value
Proposed IA	<b><math>7.43 \times 10^4</math></b>	$4.72 \times 10^3$	<b>0.0015</b>
Standard IA	$8.62 \times 10^4$	$6.37 \times 10^3$	

## Registration Results



(a) Direct registration

(b) Standard IA

(c) Proposed IA

Method	mean	std
Proposed IA	<b>83.30</b>	54.31
Standard IA	140.40	135.07
Direct registration	176.20	116.29

unit is nm, pixel size is 40nm

Tian Cao, Marc Niethammer and etc. Multi-modal registration for correlative microscopy using image analogies. Medical image analysis, 2013

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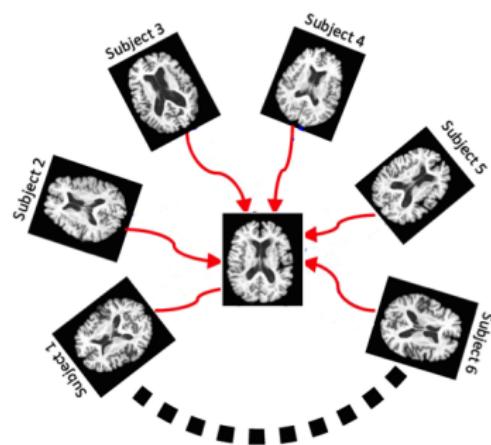
# Deformation Estimation

## Challenges

- difficult to model the relationship between deformations and appearances
- whether a basis for deformations and appearances exist

## Previous work

- Global linear regression [C.Chou2010]
- Estimate deformations of a set of key points [Q.Wang2012]

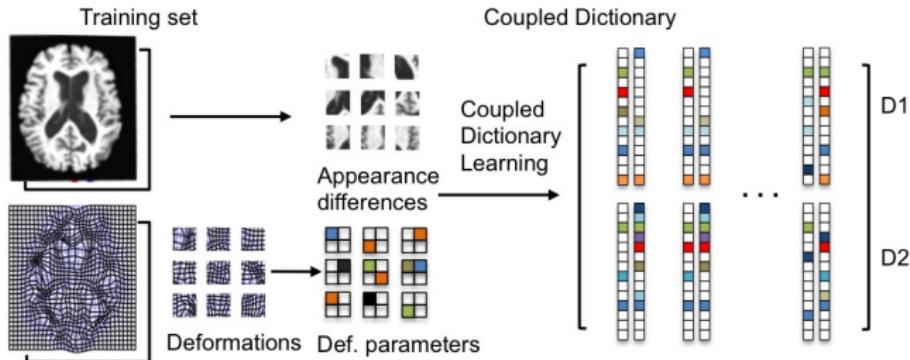


## Proposed Method

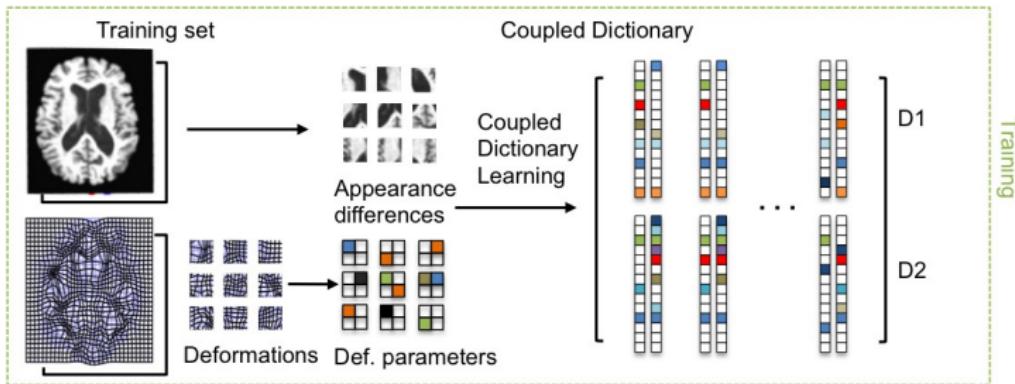
A general framework for deformation estimation based on

- semi-coupled dictionary learning on **deformation parameters and image appearances**
- investigation of different deformation parametrization methods
  - b-spline
  - initial momentum

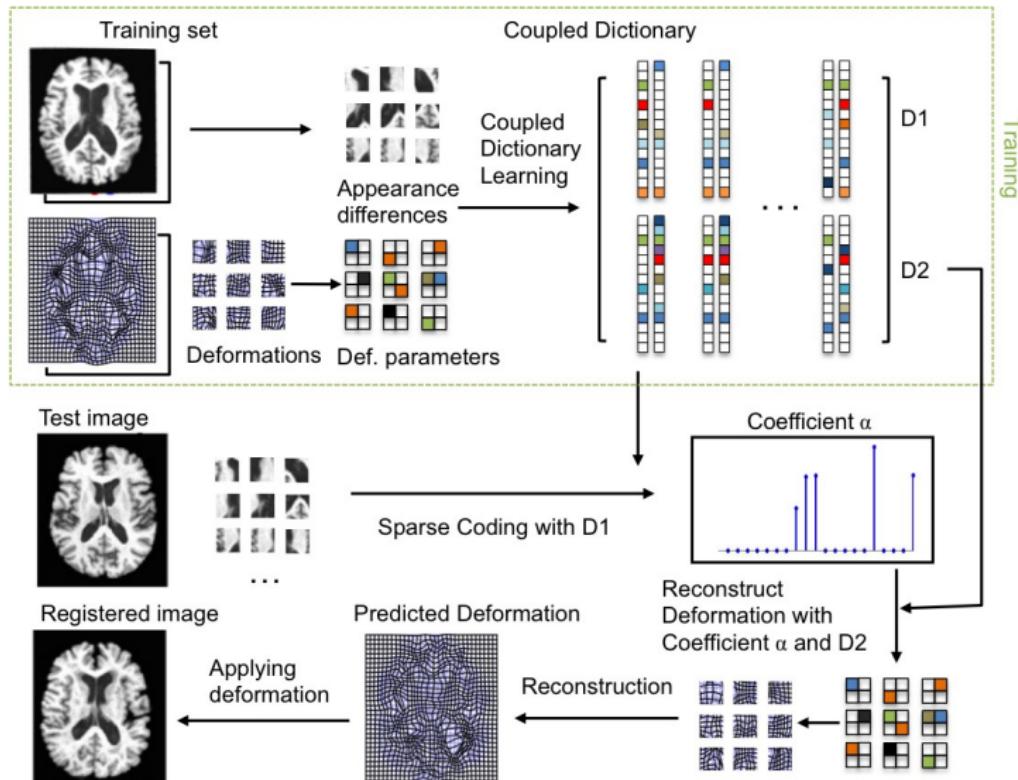
# Framework of Proposed Method



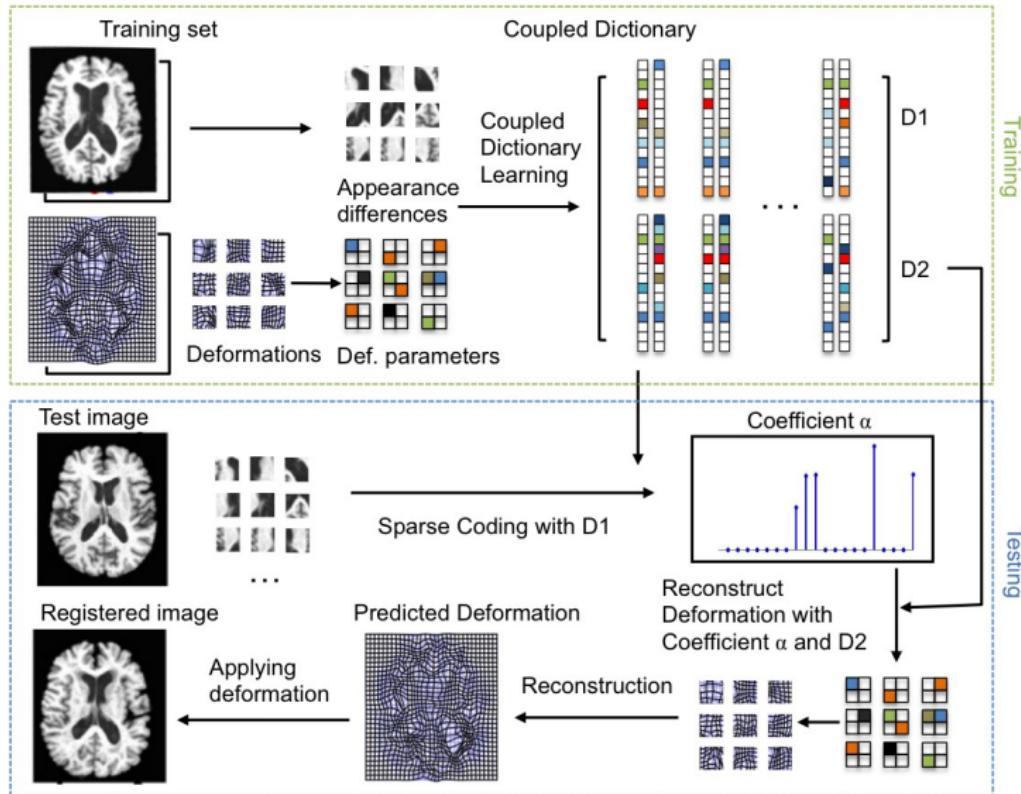
# Framework of Proposed Method



# Framework of Proposed Method



# Framework of Proposed Method



# Semi-coupled Dictionary Learning

- Dictionary learning for  $D^1$

$$\underset{D^1, D^2, \alpha_i^1, \alpha_i^2, W}{\text{minimize}} \sum_{i=1}^N \left[ \frac{1}{2} \|x_i^1 - D^1 \alpha_i^1\|_2^2 + \lambda_1 \|\alpha_i^1\|_1 \right] +$$

$$\frac{1}{2} \|x_i^2 - D^2 \alpha_i^2\|_2^2 + \lambda_2 \|\alpha_i^2\|_1 + \gamma_1 \|\alpha_i^2 - W \alpha_i^1\|_2^2 + \gamma_2 \|W\|_F^2$$

## Semi-coupled Dictionary Learning

- Dictionary learning for  $D^1$

$$\underset{D^1, D^2, \alpha_i^1, \alpha_i^2, W}{\text{minimize}} \sum_{i=1}^N \left[ \frac{1}{2} \|x_i^1 - D^1 \alpha_i^1\|_2^2 + \lambda_1 \|\alpha_i^1\|_1 \right] +$$

$$\left. \begin{aligned} & \frac{1}{2} \|x_i^2 - D^2 \alpha_i^2\|_2^2 + \lambda_2 \|\alpha_i^2\|_1 \\ & \gamma_1 \|\alpha_i^2 - W \alpha_i^1\|_2^2 + \gamma_2 \|W\|_F^2 \end{aligned} \right)$$

- Dictionary learning for  $D^2$

## Semi-coupled Dictionary Learning

- Dictionary learning for  $D^1$

$$\underset{D^1, D^2, \alpha_i^1, \alpha_i^2, W}{\text{minimize}} \sum_{i=1}^N \left[ \frac{1}{2} \|x_i^1 - D^1 \alpha_i^1\|_2^2 + \lambda_1 \|\alpha_i^1\|_1 \right] +$$

$$\frac{1}{2} \|x_i^2 - D^2 \alpha_i^2\|_2^2 + \lambda_2 \|\alpha_i^2\|_1 + \gamma_1 \|\alpha_i^2 - W \alpha_i^1\|_2^2 + \gamma_2 \|W\|_F^2$$

- Dictionary learning for  $D^2$
- Linear mapping between  $\alpha^1$  and  $\alpha^2$

## Semi-coupled Dictionary Learning

- Dictionary learning for  $D^1$

$$\underset{D^1, D^2, \alpha_i^1, \alpha_i^2, W}{\text{minimize}} \sum_{i=1}^N \left[ \frac{1}{2} \|x_i^1 - D^1 \alpha_i^1\|_2^2 + \lambda_1 \|\alpha_i^1\|_1 \right] +$$

$$\frac{1}{2} \|x_i^2 - D^2 \alpha_i^2\|_2^2 + \lambda_2 \|\alpha_i^2\|_1 + \gamma_1 \|\alpha_i^2 - W \alpha_i^1\|_2^2 + \gamma_2 \|W\|_F^2$$

- Dictionary learning for  $D^2$
- Linear mapping between  $\alpha^1$  and  $\alpha^2$
- Atoms satisfy  $\|d_i^1\|_2 \leq 1$  and  $\|d_i^2\|_2 \leq 1$

## Semi-coupled Dictionary Learning

- Dictionary learning for  $D^1$

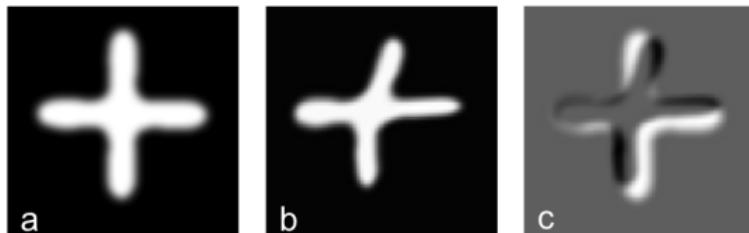
$$\underset{D^1, D^2, \alpha_i^1, \alpha_i^2, W}{\text{minimize}} \sum_{i=1}^N \left[ \frac{1}{2} \|x_i^1 - D^1 \alpha_i^1\|_2^2 + \lambda_1 \|\alpha_i^1\|_1 \right] +$$

$$\left[ \frac{1}{2} \|x_i^2 - D^2 \alpha_i^2\|_2^2 + \lambda_2 \|\alpha_i^2\|_1 \right] + \gamma_1 \|\alpha_i^2 - W \alpha_i^1\|_2^2 + \gamma_2 \|W\|_F^2$$

- Dictionary learning for  $D^2$
- Linear mapping between  $\alpha^1$  and  $\alpha^2$
- Atoms satisfy  $\|d_i^1\|_2 \leq 1$  and  $\|d_i^2\|_2 \leq 1$
- Use a linear mapping  $W$  to relating  $\alpha^1$  and  $\alpha^2$  to relax the strong coupling

## Results

- synthesize cross image as atlas
- randomly generate  $3 \times 3$  b-spline coefficients to transform atlas as training subject images
- test on 100 images with random b-spline transformations



## Results

Method	median	mean	min	max	std
RAW	17.9776	19.2912	6.4132	45.1841	9.1304
NN	6.4063	7.1153	1.1399	12.2234	4.0948
GR	4.9261	5.2245	2.6279	10.2138	3.7925
CDL	3.6325	3.7531	0.724	8.5439	2.1158
SCDL	<b>3.0679</b>	<b>2.9254</b>	<b>0.3532</b>	<b>5.0559</b>	<b>1.4815</b>

Unit: pixel

RAW: direct registration

NN: nearest neighbor search

GR: global regression

CDL: coupled dictionary learning

SCDL: semi-coupled dictionary learning

mean absolute errors (MAE) in pixels

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Tian Cao, Nikhil Singh, Vladimir Jojic, and Marc Niethammer.  
Semi-coupled dictionary learning for deformation prediction. Accepted by ISBI  
2015.

# Outline

- Background
- CDL for Multi-modal Image registration
- CDL for Deformation Estimation
- CDL for GTPase Activations and Cell Protrusions
- Robust CDL
- Summary and Future Work

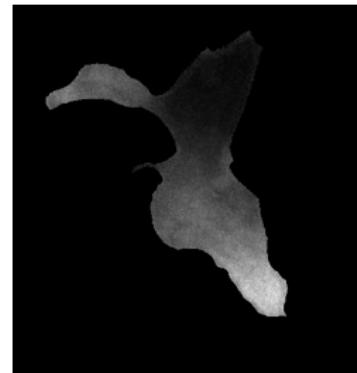
# GTPase Activations and Cell Protrusions Modeling

## Challenges

- how to represent the data
- how to model the global and local relationships

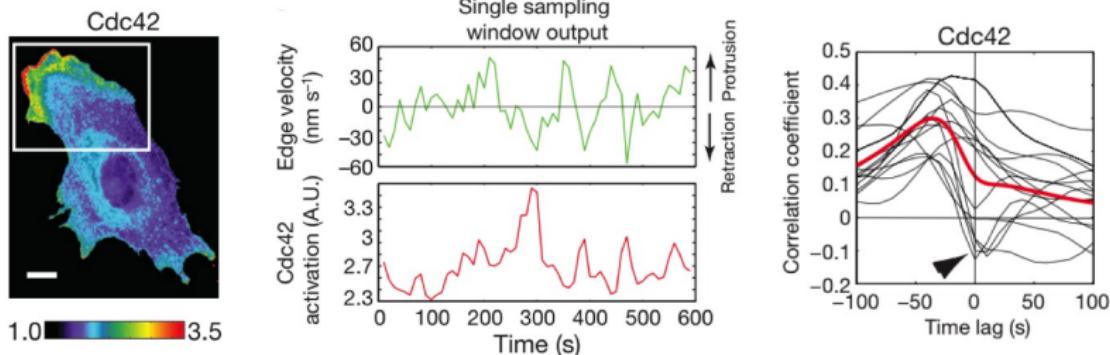
## Previous work

- compute correlation between activations and protrusions on boundary points [M. Machacek 2009]



## Previous Work

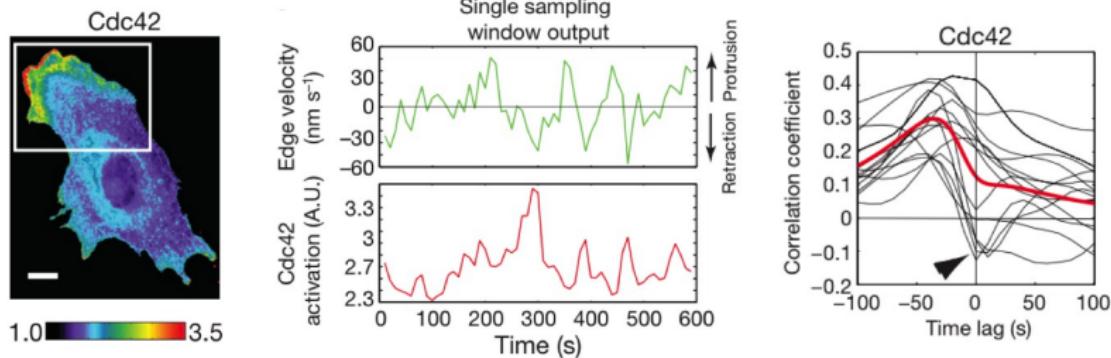
- Track boundary points in specific regions
- Record intensity values for small windows centered at boundary points
- Compute cross-correlation between GTPase activations and cell movements on boundary points



[Matthias Machacek and etc, Nature letter, 2009]

# Limitations of Previous Work

- regions of interest are hand-picked
- missing spatial interaction between neighboring boundary points



[Matthias Machacek and etc, Nature letter, 2009]

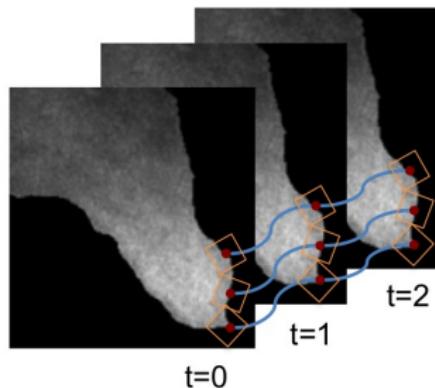
## Proposed Method

A framework to model the GTPase activations and cell movements based on

- coupled dictionary learning for **spatiotemporal data**
  - GTPase activations and cell movements on boundary points
- represent data with sparse coefficients
- clustering boundary points based on sparse coefficients

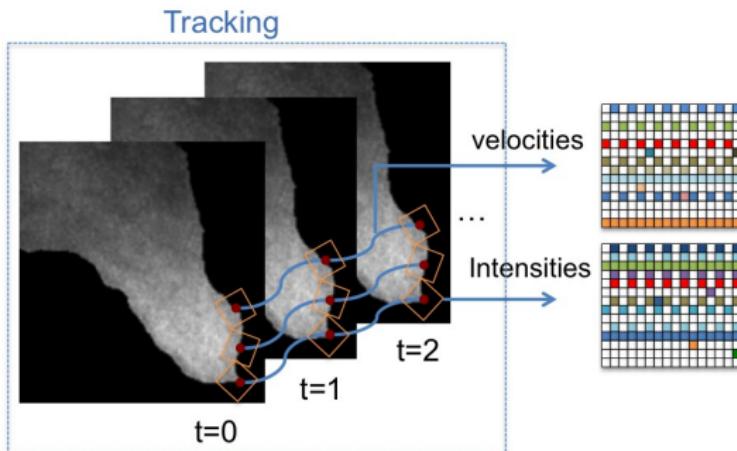
## Proposed Method

- Learn coupled dictionary on protrusion and activation for single boundary points



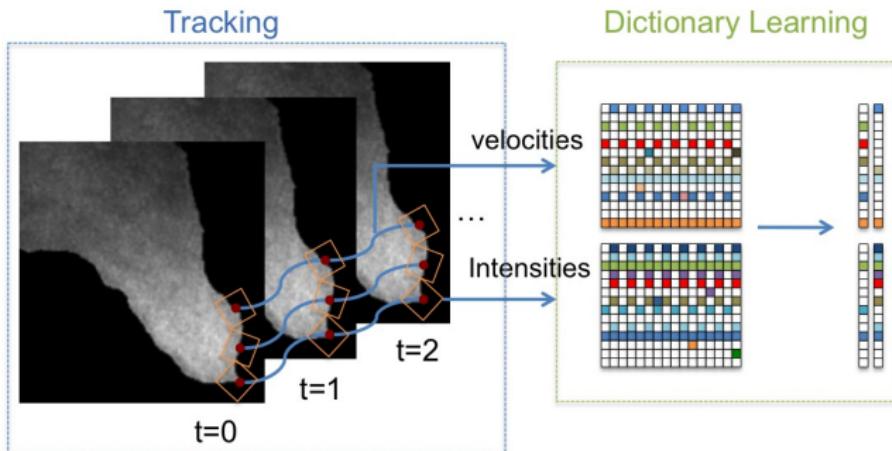
# Proposed Method

- Learn coupled dictionary on protrusion and activation for single boundary points



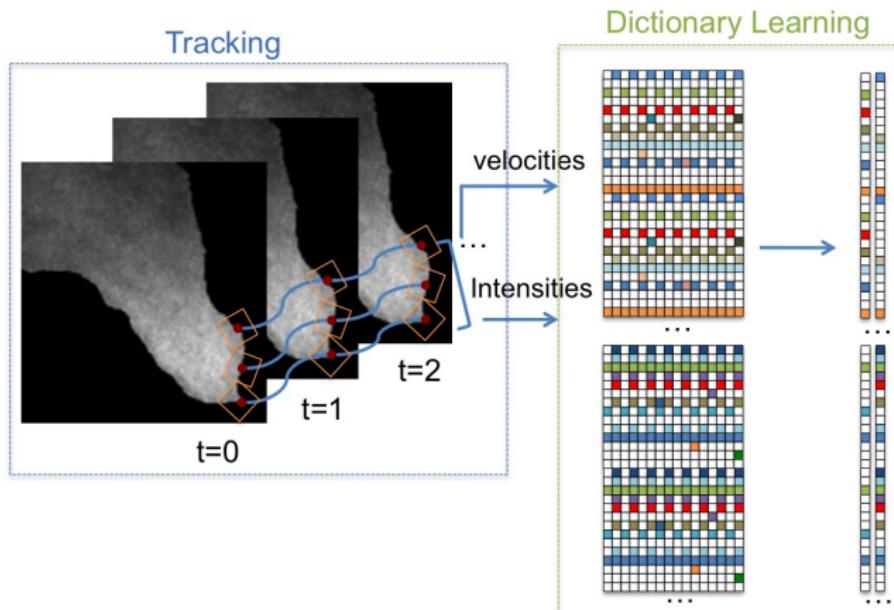
# Proposed Method

- Learn coupled dictionary on protrusion and activation for single boundary points



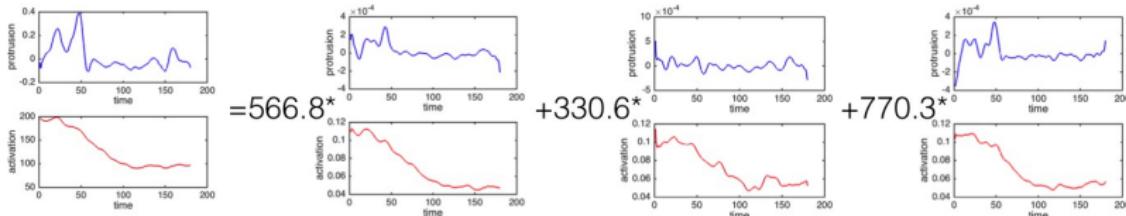
# Proposed Method

- Learn coupled dictionary on protrusion and activation for single boundary points
- Learn coupled dictionary on protrusion and activation for boundary point “patches”



# Proposed Method

- Use coupled dictionary to reconstruct the data
- Represent data with compact coefficients based on learned dictionary

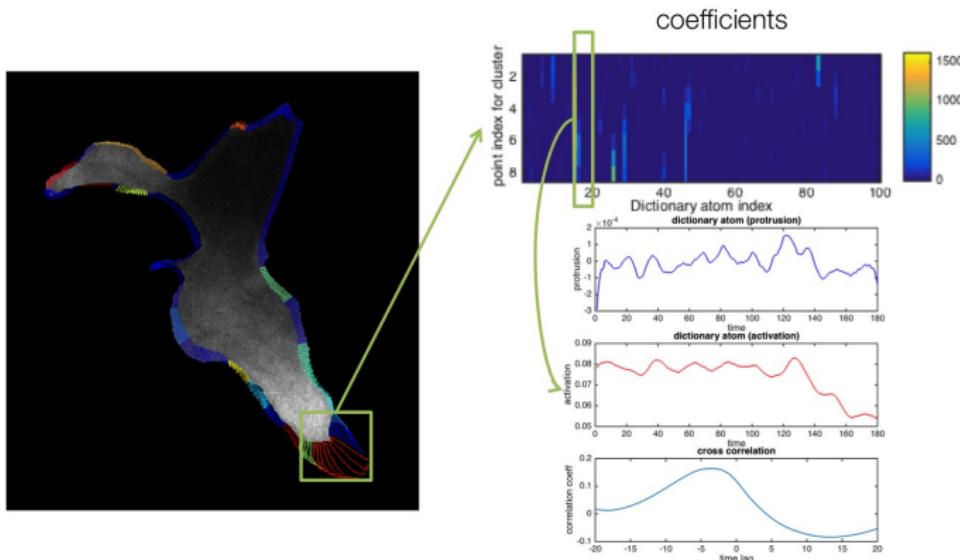


$$\text{Coefficients} = [0, 0, \dots, 566.8, 0, 0, \dots, 330.6, 0, 0, \dots, 770.3]$$

# Preliminary Results

To study spatial interactions between neighboring points or “patches”

- Represent data with compact coefficients based on learned dictionary
- k-means clustering on coefficients of points or “patches”



## Hierarchical clustering

### Limitations of K-means clustering

- need assign number of clusters in k-means clustering
- find optimal k is difficult

### Hierarchical clustering

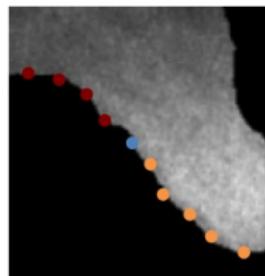
- arrange the data in a tree structure
- provide hierarchical relationship between different data points

## Discussion

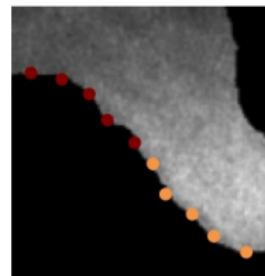
Why do clustering on coefficients instead of raw data?

- coefficients can provide information about the patterns used to represent the data

Why do clustering on “point patches”?



clustering on single points



clustering on point “patches”

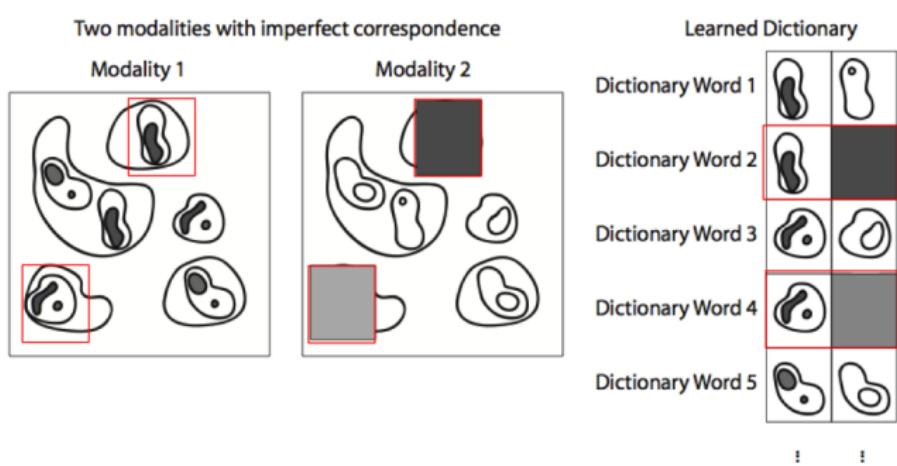
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# Robust Coupled Dictionary Learning

## Challenges for Coupled dictionary learning

- may fail without sufficient correspondences between training data in two spaces



## Our Method

A probabilistic model for coupled dictionary learning

- confidence measure
  - a conditional probability  $p(h|x_i)$
  - $h = 0$  indicates  $x_i$  reconstructed by “clean” dictionary atoms
  - $h = 1$  indicated  $x_i$  reconstructed by “noise” dictionary atoms
- EM algorithm

## Confidence Measure

- Applying Bayes' rules,  $p(h = 1|x_i)$  can be represented as

$$P(h = 1|x_i) = \frac{P(x_i|h = 1)P(h = 1)}{P(x_i|h = 1)P(h = 1) + P(x_i|h = 0)P(h = 0)}$$

- Assuming independence of  $x_i$  and the Gaussian distribution of pixels in  $x_i$

$$p(x_i|h = 1, \theta_1) = \mathcal{N}(x_i; \mu_1, \sigma_1^2),$$

$$p(x_i|h = 0, \theta_0; D, \alpha_i) = \mathcal{N}(x_i - D\alpha_i; 0, \sigma_0^2).$$

## EM Algorithm

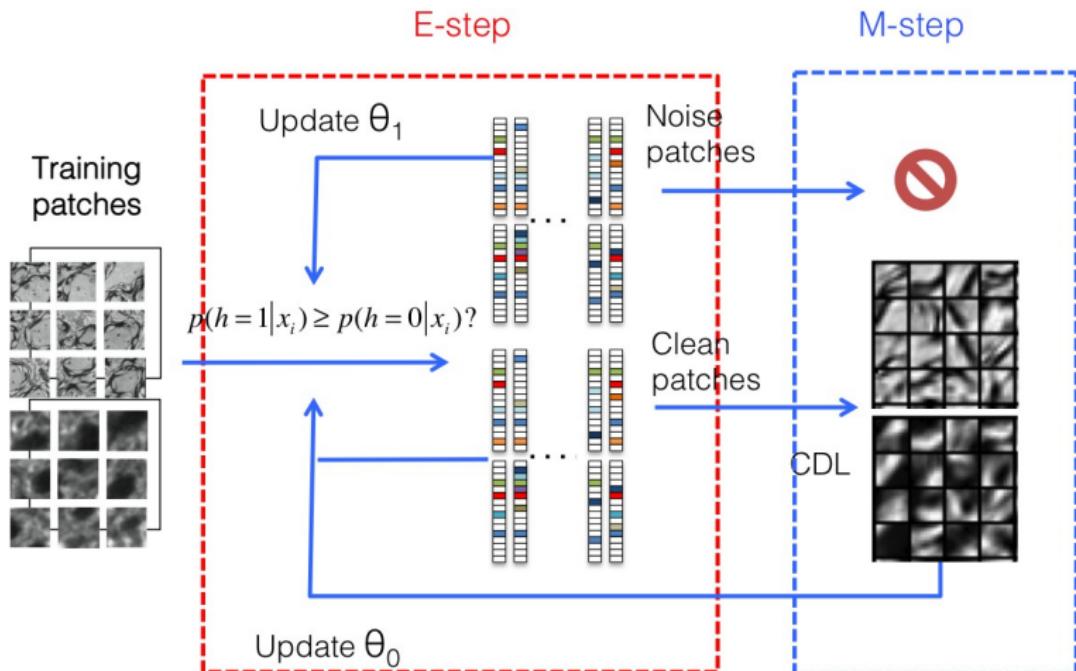
The parameters to estimate are

- $\theta_1 = \{\mu_i, \sigma_1\}$
- $\theta_o = \sigma_0$

Maximum likelihood (ML) estimation

- classify non-corresponding data and update coupled dictionary iteratively

# Framework of Proposed Method



## Results

	Training images	Learned dictionaries	
		Standard method	Proposed method
Modality 1			

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Tian Cao, Vladimir Jovic, Marc Niethammer and etc. Robust multi-modal dictionary learning. MICCAI, 2013.

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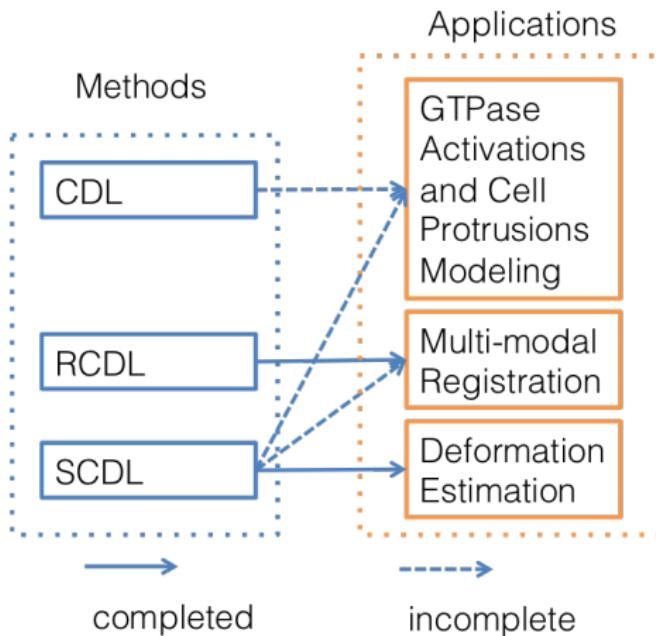
## Summary

Coupled dictionary learning method to relate data from two spaces:

- spatial data: appearances for different modalities
- spatial data: appearances and deformations
- spatiotemporal data: GTPase activations and cell movements

A probabilistic model for robust coupled dictionary learning to account for data non-corresponding training data

# Summary and Future Work



## Research Status and Tentative Timeline

CDL for multi-modal registration	Completed([1, 4])
CDL for deformation prediction	Completed([3])
CDL for GTPase activation and cell protrusion modeling	May 2015
Robust CDL	Completed([2])
Future Work	June 2015
Proposal	January 2015
Oral Exam	February 2015
Defense	August 2015

Table: Research plan

## List of publications

-  T. Cao, C. Zach, S. Modla, D. Powell, K. Czummek, and M. Niethammer. [Registration for correlative microscopy using image analogies](#). *Biomedical Image Registration*, pages 296–306, 2012.
-  Tian Cao, Vladimir Jojic, Shannon Modla, Debbie Powell, Kirk Czummek, and Marc Niethammer. [Robust multimodal dictionary learning](#). In *Medical Image Computing and Computer-Assisted Intervention–MICCAI 2013*, pages 259–266. Springer Berlin Heidelberg, 2013.
-  Tian Cao, Nikhil Singh, Vladimir Jojic, and Marc Niethammer. [Semi-coupled dictionary learning for deformation prediction](#). In *Accepted by ISBI 2015*.
-  Tian Cao, Christopher Zach, Shannon Modla, Debbie Powell, Kirk Czummek, and Marc Niethammer. [Multi-modal registration for correlative microscopy using image analogies](#). *Medical image analysis*, 2013.

# Thesis

- Learning a coupled basis for the compact representation of two spaces can be achieved by **coupled dictionary learning** (CDL).
- Such dictionaries can be learned to capture appearance differences of different imaging modalities, dependencies between image appearance and deformation as well as the spatiotemporal patterns for cell signaling and boundary protrusions and retractions.
- To account for data inconsistencies, a robust coupled dictionary can be obtained based on a probabilistic dictionary model.