

# Simultaneous Sparsity Model for Histopathological Image Classification

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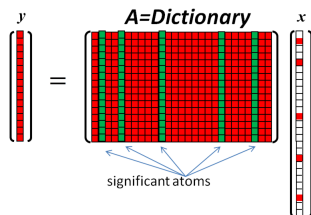
<sup>3</sup>Dept. of Veterinary and Biomedical Sciences, Pennsylvania State University



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# The compressed sensing (CS) problem<sup>2</sup>



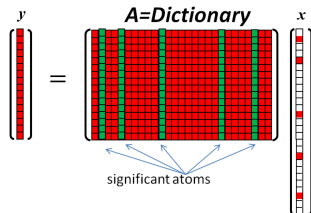
How to represent  $y$  with the **fewest** number of dictionary atoms?

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<sup>1</sup>Donoho, Communications on Pure and Applied Mathematics, 2006

<sup>2</sup>Candes et al., IEEE Transactions on Information Theory, 2006

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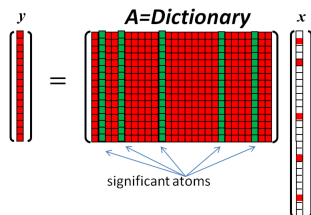
$$\min_{\mathbf{x}} \|\mathbf{x}\|_0 \text{ subject to } \mathbf{y} = \mathbf{A}\mathbf{x}$$

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Convex relaxation<sup>1</sup>:

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# Sparse representation-based image classification<sup>3</sup>

- Extension of CS analytical framework to classification by designing **class-specific** dictionaries

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## Sparse representation-based image classification<sup>3</sup>

- Extension of CS analytical framework to classification by designing **class-specific** dictionaries
- Given:  $M$  classes, with  $N_i$  training images in the  $i$ -th class
- **Assumption:** New image from class  $i$  lies (approximately) in the linear span of training samples from class  $i$

$$\mathbf{y} = x_{i,1}\mathbf{a}_{i,1} + x_{i,2}\mathbf{a}_{i,2} + \dots + x_{i,N_i}\mathbf{a}_{i,N_i} = \mathbf{A}_i\mathbf{x}_i$$

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- $\mathbf{y} \rightarrow$  **sparse** linear combination of **all** training samples:

$$\mathbf{y} = \begin{bmatrix} \mathbf{A}_1 & \mathbf{A}_2 & \dots & \mathbf{A}_M \end{bmatrix} \begin{bmatrix} \mathbf{x}_1 \\ \mathbf{x}_2 \\ \vdots \\ \mathbf{x}_M \end{bmatrix} = \mathbf{A}\mathbf{x}$$

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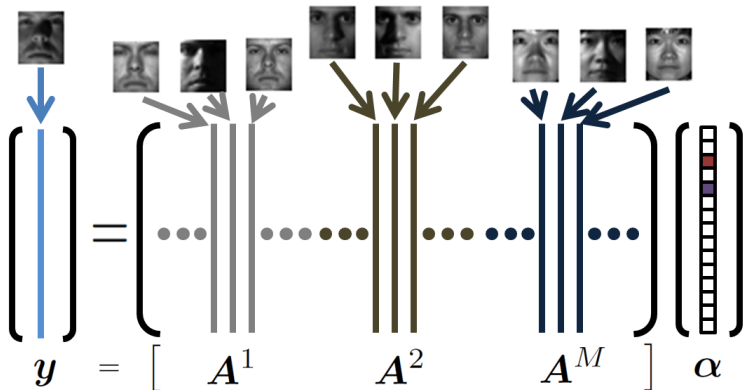
- Membership of  $\mathbf{y}$  encoded by sparse representation

$$\mathbf{x} = [\mathbf{0}^t \quad \dots \quad \mathbf{0}^t \quad \mathbf{x}_i^t \quad \mathbf{0}^t \quad \dots \quad \mathbf{0}^t]^t.$$

<sup>3</sup>Wright et al., IEEE Transactions on Pattern Analysis and Machine Intelligence, 2009



# Application: Face recognition



# Sparse representation-based classification (SRC)

- Solve the sparse recovery problem:

$$\hat{\mathbf{x}} = \arg \min \|\mathbf{x}\|_1 \quad \text{subject to} \quad \|\mathbf{y} - \mathbf{Ax}\|_2 \leq \epsilon$$

# Sparse representation-based classification (SRC)

- Solve the sparse recovery problem:

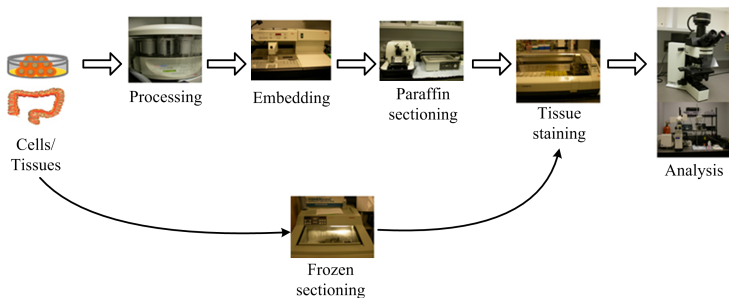
$$\hat{\mathbf{x}} = \arg \min \|\mathbf{x}\|_1 \quad \text{subject to} \quad \|\mathbf{y} - \mathbf{A}\mathbf{x}\|_2 \leq \epsilon$$

- Class decision based on reconstruction residuals:

$$\text{identity}(\mathbf{y}) = \arg \min_i \|\mathbf{y} - \mathbf{A}\delta_i(\hat{\mathbf{x}})\|_2$$

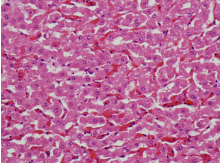
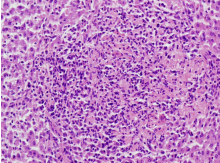
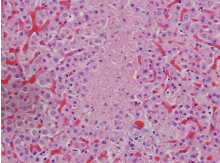
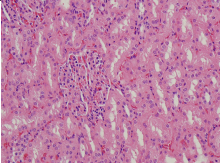
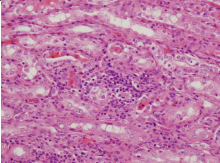
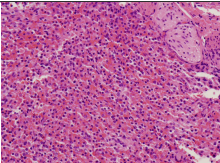
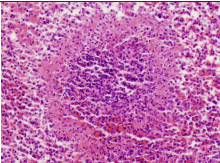
$\delta_i(\hat{\mathbf{x}}) \rightarrow$  only non-zero entries are those associated with class  $i$

# Histopathology



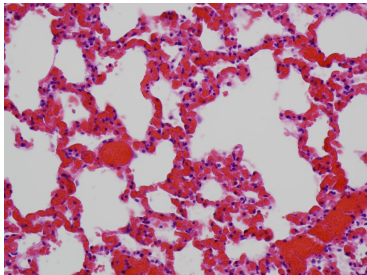
- High resolution images → underlying **tissue architecture** preserved
- H&E-staining
  - Hematoxylin → bluish nuclei
  - Eosin → red cytoplasm and connective tissue
- **Applications:** cancer or more generally **disease detection and grading** → breast, prostate, lung, liver, spleen, etc.

# Example histopathological images<sup>4</sup>

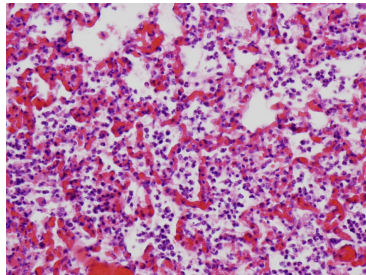
Organ	Healthy	Inflammation	Necrosis
Liver			
Kidney			-
Spleen			-

<sup>4</sup>Animal Diagnostics Laboratory, Penn State

## Example images: Lung



(a) Healthy lung.



(b) Lung inflammation.

# Opportunities and challenges

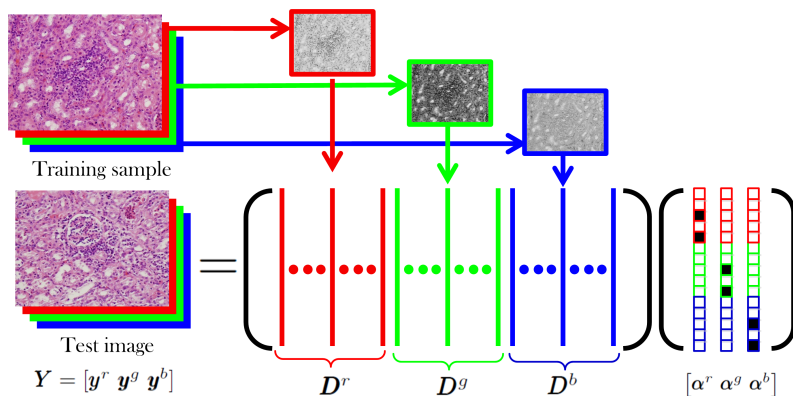
- Multi-channel (color) information
  - Extension of sparsity-based techniques via color dictionaries
  - Maintain color channel correspondence

# Opportunities and challenges

- 1 Multi-channel (color) information
  - Extension of sparsity-based techniques via color dictionaries
  - Maintain color channel correspondence
- 2 Rich structural information in tissues
  - Feature extraction is a challenging task → customized to specific organs
  - Example-based learning



# SHIRC: Simultaneous Sparsity model for Histopathological Image Representation and Classification



# Designing color channel constraints

$$\mathbf{Y} = \mathbf{DS} = \begin{bmatrix} \mathbf{D}_h^r & \mathbf{D}_d^r & \mathbf{D}_h^g & \mathbf{D}_d^g & \mathbf{D}_h^b & \mathbf{D}_d^b \end{bmatrix} \begin{bmatrix} \boldsymbol{\alpha}^r & \boldsymbol{\alpha}^g & \boldsymbol{\alpha}^b \end{bmatrix}$$

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$$\alpha^r = \begin{bmatrix} \alpha_h^r \\ \alpha_d^r \\ 0 \\ 0 \\ 0 \\ 0 \end{bmatrix}, \alpha^g = \begin{bmatrix} 0 \\ 0 \\ \alpha_h^g \\ \alpha_d^g \\ 0 \\ 0 \end{bmatrix}, \alpha^b = \begin{bmatrix} 0 \\ 0 \\ 0 \\ 0 \\ \alpha_h^b \\ \alpha_d^b \end{bmatrix}$$

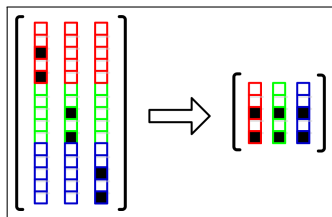
- $\alpha^r, \alpha^g, \alpha^b \in \mathbb{R}^{3n}$

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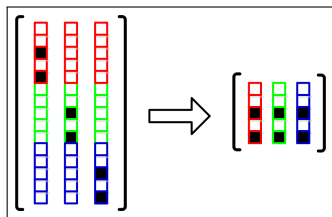


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$$S' = \begin{bmatrix} \alpha_h^r & \alpha_h^g & \alpha_h^b \\ \alpha_d^r & \alpha_d^g & \alpha_d^b \end{bmatrix}$$

$$S \in \mathbb{R}^{3n \times 3} \xrightarrow{?} S' \in \mathbb{R}^{n \times 3}$$

# Formulating the optimization problem

Define:

$$\mathbf{H} = \begin{bmatrix} \mathbf{1}_n & \mathbf{0}_n & \mathbf{0}_n \\ \mathbf{0}_n & \mathbf{1}_n & \mathbf{0}_n \\ \mathbf{0}_n & \mathbf{0}_n & \mathbf{1}_n \end{bmatrix} \in \mathbb{R}^{3n \times 3n}, \quad \mathbf{J} = [\mathbf{I}_n \quad \mathbf{I}_n \quad \mathbf{I}_n] \in \mathbb{R}^{n \times 3n}$$

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Hadamard operator:

$$\mathbf{H} \circ \mathbf{S} = \begin{bmatrix} \boldsymbol{\alpha}^r & \mathbf{0} & \mathbf{0} \\ \mathbf{0} & \boldsymbol{\alpha}^g & \mathbf{0} \\ \mathbf{0} & \mathbf{0} & \boldsymbol{\alpha}^b \end{bmatrix} \in \mathbb{R}^{3n \times 3n}$$

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$$\mathbf{J}(\mathbf{H} \circ \mathbf{S}) = [\mathbf{I}_n \quad \mathbf{I}_n \quad \mathbf{I}_n] \begin{bmatrix} \boldsymbol{\alpha}^r & \mathbf{0} & \mathbf{0} \\ \mathbf{0} & \boldsymbol{\alpha}^g & \mathbf{0} \\ \mathbf{0} & \mathbf{0} & \boldsymbol{\alpha}^b \end{bmatrix} = [\boldsymbol{\alpha}^r \quad \boldsymbol{\alpha}^g \quad \boldsymbol{\alpha}^b] = \mathbf{S}'$$



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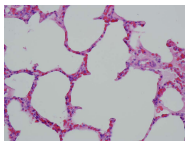
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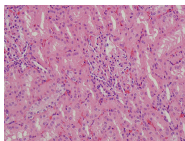
$$\hat{\mathbf{S}}' = \arg \min \|\mathbf{S}'\|_{\text{row},0} \quad \text{subject to} \quad \|\mathbf{Y} - \mathbf{D}\mathbf{S}\|_F \leq \epsilon$$

# Experimental set-up: Data sets

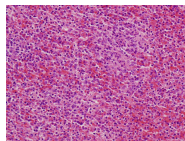
- ADL data set<sup>5</sup>
  - Mammalian tissue
  - Three organs: kidney, lung, spleen (healthy vs. inflammation)
  - 40 training images, 80 test images



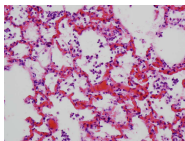
(a) Lung



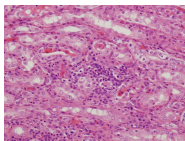
(b) Kidney



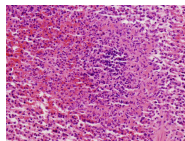
(c) Spleen



(d) Lung



(e) Kidney



(f) Spleen

Figure : Top row: healthy; bottom row: inflammatory.

<sup>5</sup> Animal Diagnostics Laboratory, Penn State

# Experimental set-up: Data sets

- IBL data set<sup>6</sup>
  - Human intraductal breast lesions
  - Ductal carcinoma in situ (DCIS) vs. usual ductal hyperplasia (UDH)
  - 32 training images, 22 test images

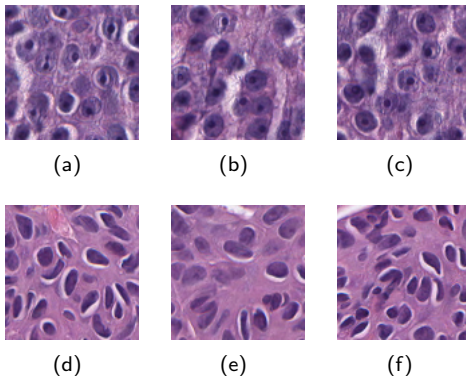


Figure : Top row: DCIS; bottom row: UDH.

<sup>6</sup>Clarian Pathology Lab, Indianapolis; Computer and Information Science Dept., IUPUI

# Experimental set-up

## Comparison with:

- 1 SRC: using luminance channel information only
- 2 SVM: State-of-the-art WND-CHARM features<sup>7,8</sup>
  - Image pixel statistics
  - Color channel histograms
  - Gray level co-occurrence matrix (GLCM)
  - Wavelet coefficients
  - Morphological features

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<sup>7</sup> Orlov et al., Pattern Recognition Letters, 2008

<sup>8</sup> <http://ome.grc.nia.nig.gov/wnd-charm>

# Results I: Confusion matrix

- Each row: true class identity of image
- Each column: class predicted by the classifier

Table : Lung (ADL)

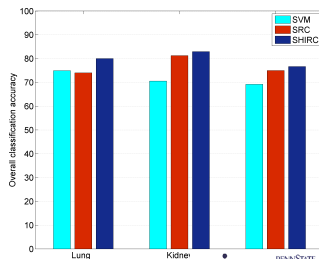
Class	Healthy	Inflammatory	Method
Healthy	<b>0.8875</b>	0.1125	SVM
	0.7250	0.2750	SRC
	0.7500	0.2500	SHIRC
Inflammatory	0.3762	0.6238	SVM
	0.2417	0.7583	SRC
	0.1500	<b>0.8500</b>	SHIRC

Table : Kidney (ADL)

Class	Healthy	Inflammatory	Method
Healthy	0.6925	0.3075	SVM
	<b>0.8750</b>	0.1250	SRC
	0.8250	0.1750	SHIRC
Inflammatory	0.2812	0.7188	SVM
	0.2500	0.7500	SRC
	0.1667	<b>0.8333</b>	SHIRC

Table : Spleen (ADL)

Class	Healthy	Inflammatory	Method
Healthy	0.5112	0.4888	SVM
	<b>0.7083</b>	0.2917	SRC
	0.6500	0.3500	SHIRC
Inflammatory	0.1275	0.8725	SVM
	0.2083	0.7917	SRC
	0.1167	<b>0.8833</b>	SHIRC



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Table : IBL<sup>a</sup>

Class	UDH	DCIS	Method
UDH	0.7140	0.2860	SRC
	<b>0.8540</b>	0.1460	SHIRC
DCIS	0.1750	0.8250	SRC
	0.1220	<b>0.8780</b>	SHIRC

<sup>a</sup>Dundar et al., IEEE Transactions on Biomedical Engineering, 2011

## Results II: ROC curves

- Miss: diseased identified as healthy
- False alarm: healthy classified as diseased

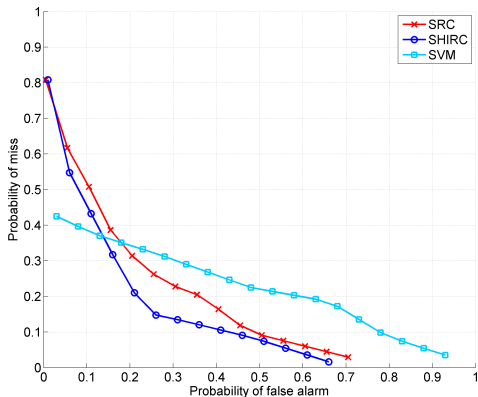


Figure : Lung (ADL).

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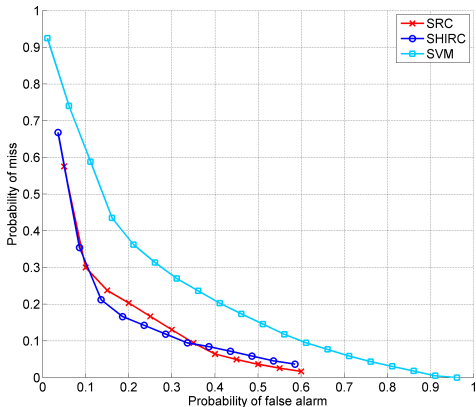


Figure : Kidney (ADL).



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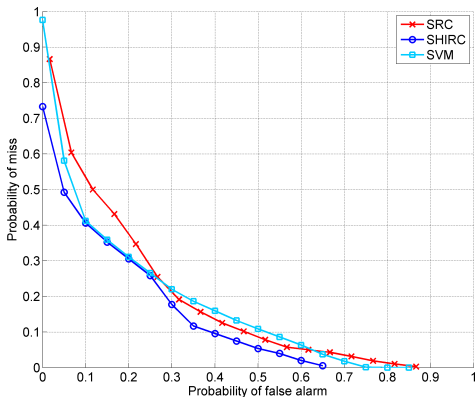


Figure : Spleen (ADL).

# Conclusions

- Joint sparsity models incorporate contextual information
  - Color channel information in medical imaging
- New greedy algorithm<sup>9</sup> for multi-dictionary sparsity-based classification

## Ongoing/Future research:

- Histopathological objects of interest (cells, nuclei) at different sizes and scales → locally adaptive framework
- Effect of training set size on classification accuracy.

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<sup>9</sup> MATLAB code: [signal.ee.psu.edu/histimg.html](http://signal.ee.psu.edu/histimg.html)

Thank you

Questions?