

Big Data Analytics for Healthcare

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Healthcare Analytics using Electronic Health Records (EHR)

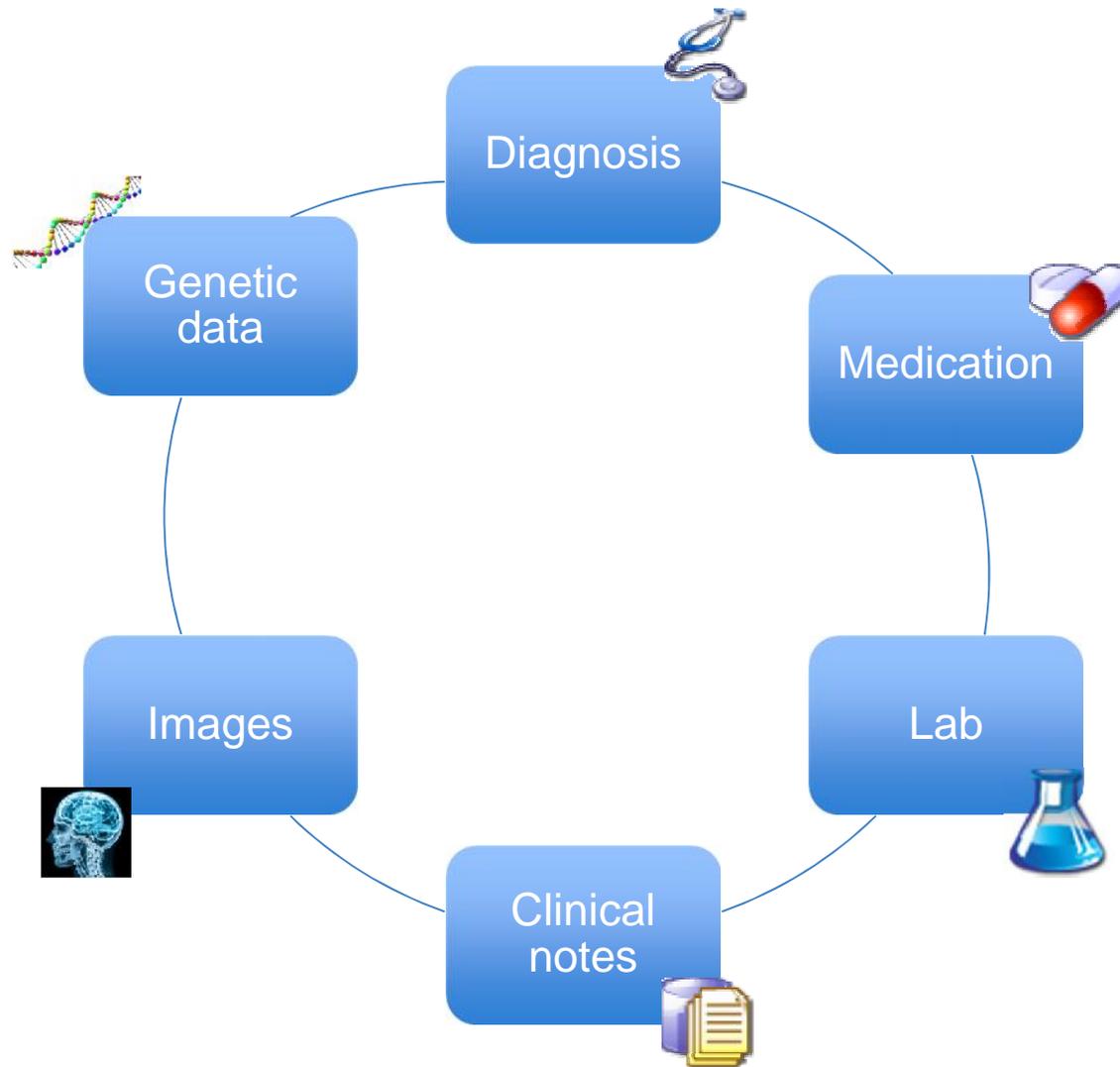


- Old way: **Data are expensive and small**
 - Input data are from clinical trials, which is small and costly
 - Modeling effort is small since the data is limited
 - A single model can still take months

- EHR era: **Data are cheap and large**
 - Broader patient population
 - Noisy data
 - Heterogeneous data
 - Diverse scale
 - Complex use cases



Heterogeneous Medical Data



Challenges in Healthcare Analytics

Collaboration across domains

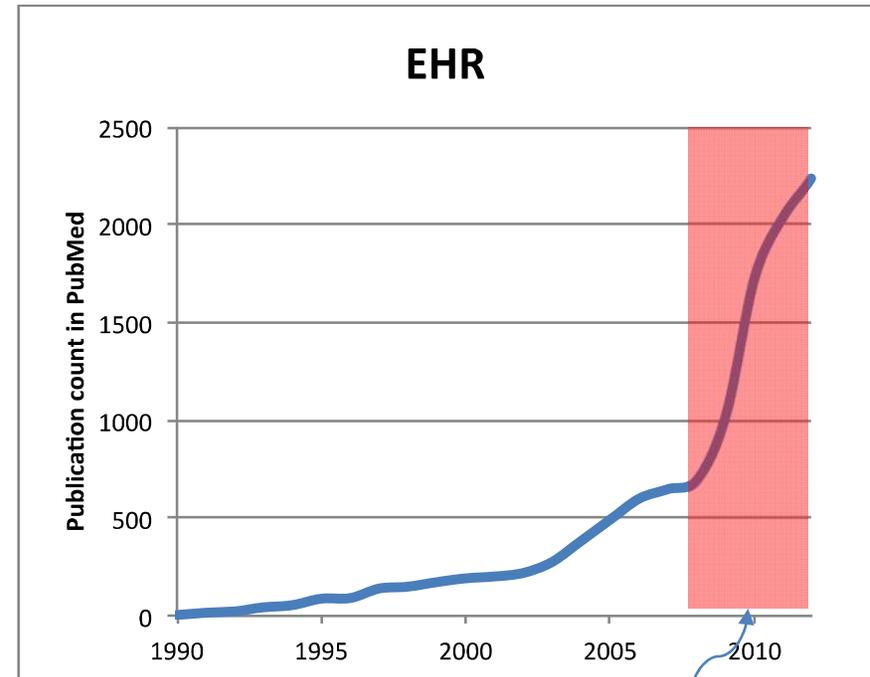
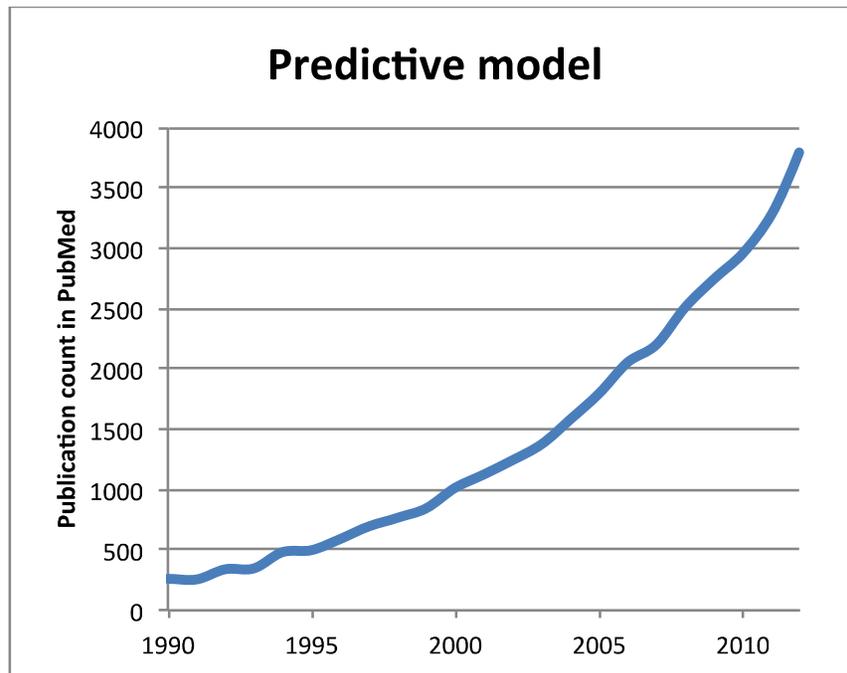
Analytic platform

Intuitive results

Scalable computation

PARALLEL MODEL BUILDING

Motivation – Predictive modeling using EHR is growing



Explosion in interest

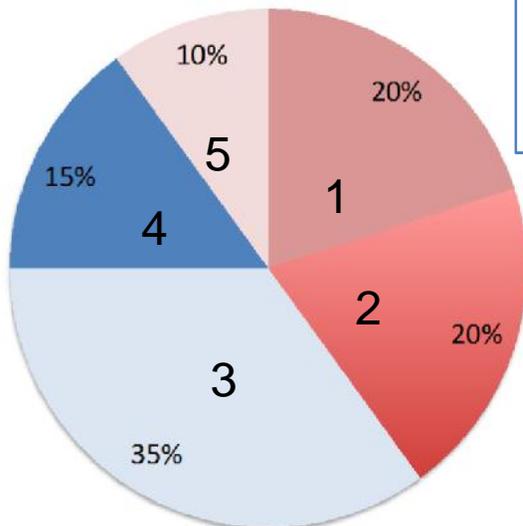
- Need for scalable predictive modeling platforms/systems due to increased computational requirements from:
 - Processing EHR data (due to volume, variability, and heterogeneity)
 - Building accurate models
 - Building clinically meaningful models
 - Validating models for accuracy and generalizability

What does it take to develop a predictive model using EHR?



Marina: IBM
Analytics Consultant

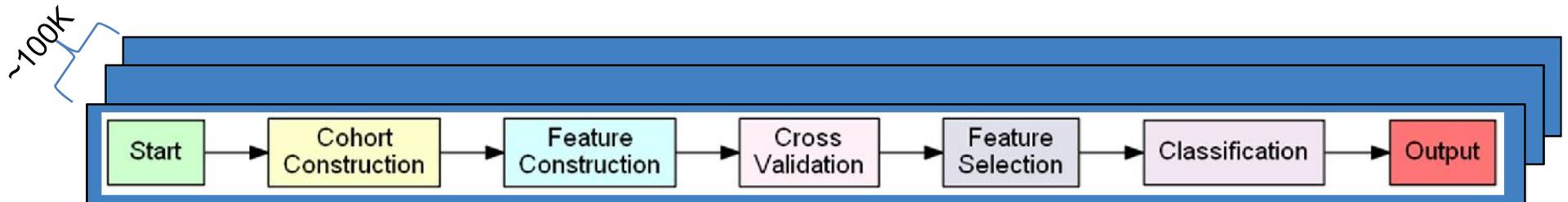
- Within 3 months, we need to
1. understand business case
 2. obtain the data
 3. prepare the data
 4. develop predictive models
 5. deliver the final model



How to help her to develop a good predictive model quickly?

David Gotz, Harry Starvropoulos, Jimeng Sun, Fei Wang.
ICDA: A Platform for Intelligent Care Delivery Analytics, AMIA 2012

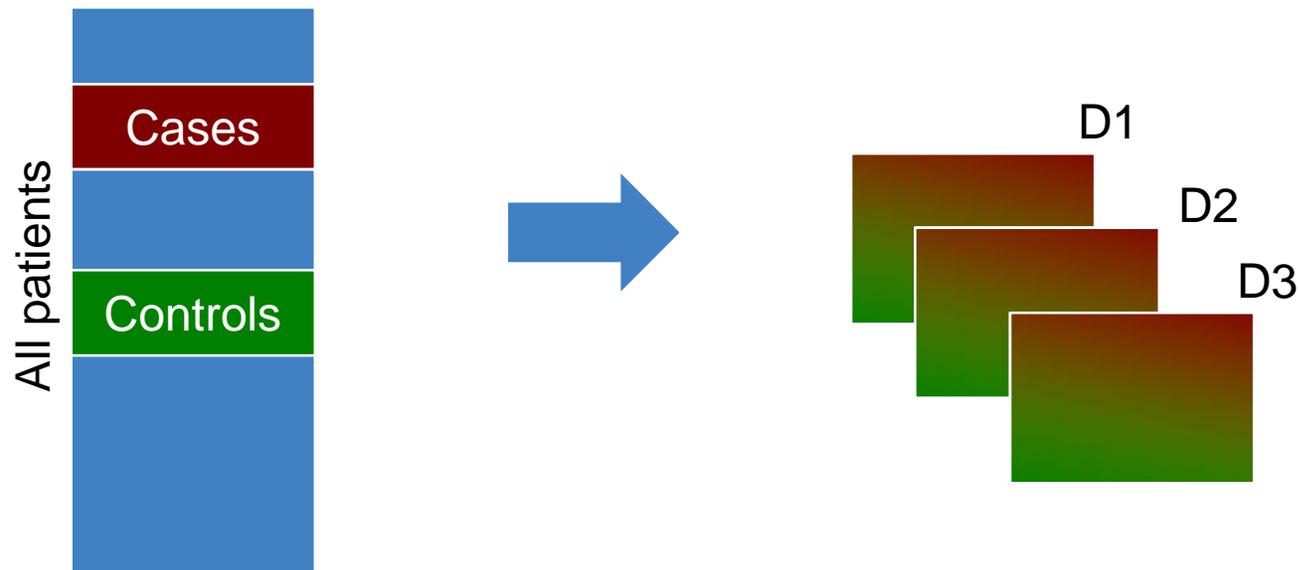
A Generalized Predictive Modeling Pipeline



Model specification

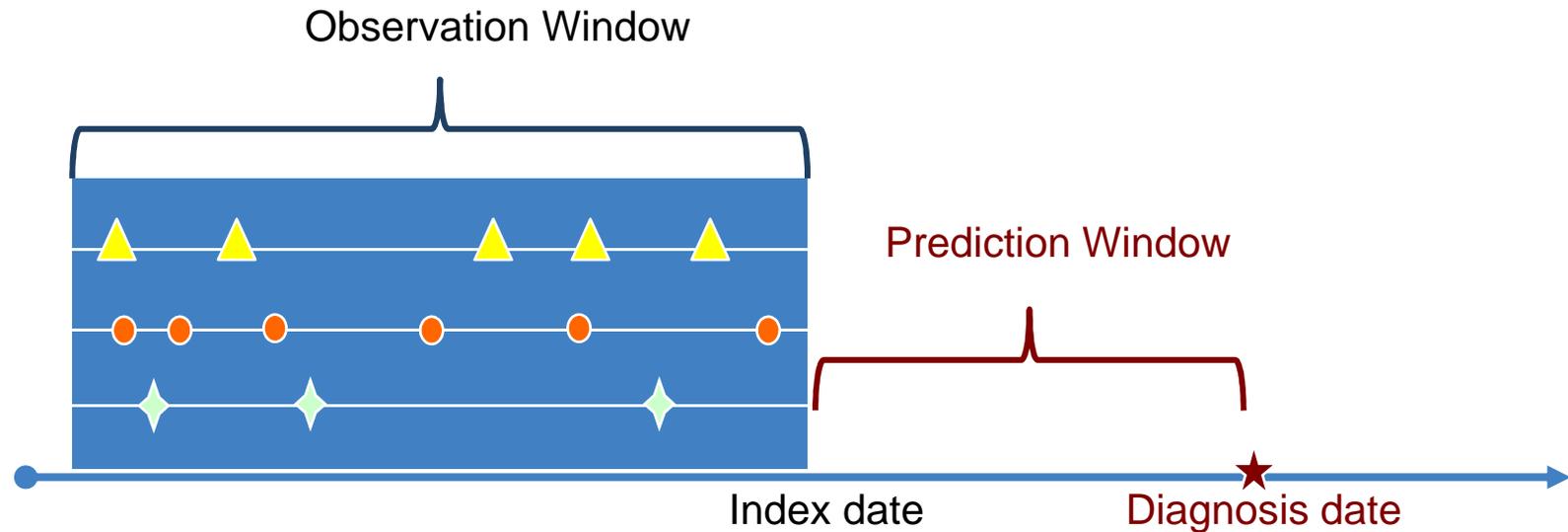
- **Cohort Construction:** Find an appropriate set of patients with the specified target condition and a corresponding set of control patients without the condition.
- **Feature Construction:** Compute a feature vector representation for each patient based on the patient's EHR data.
- **Cross Validation:** Partition the data into complementary subsets for use in model training and validation testing.
- **Feature Selection:** Rank the input features and select a subset of relevant features for use in the model.
- **Classification:** The training and evaluation of a model for a specific classifier.
- **Output:** Clean up intermediate files and to put results into their final locations.

Cohort Construction



	Disease Target	samples
D1	Hypertension control	5000
D2	Heart failure onset	33K
D3	Hypertension diagnosis	300K

Feature Construction



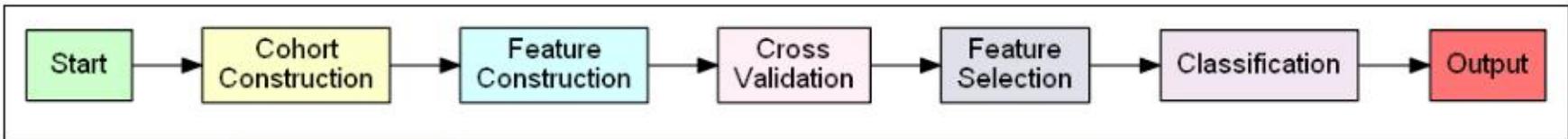
- We define
 - Diagnosis date and index date
 - Prediction and observation windows
- Features are constructed from the observation window and predict HF onset after the prediction window

PARAMO

▼ Predictive Modeling Pipeline

Pipeline: Example 1 ▾ Actions ▾

Configuration Graph Status/Results



Cohort Construction Feature Construction Cross Validation Feature Selection Classification Other

Parameter	Value
Prediction Window (days)	0
Observation Window (days)	720
Feature	Type: <input type="text" value="Diagnosis"/> Aggregation: <input type="text" value="Sum"/>
Feature	Type: <input type="text" value="Medication"/> Aggregation: <input type="text" value="Sum"/>
Feature	Type: <input type="text" value="Lab"/> Aggregation: <input type="text" value="Mean"/>
Feature	Type: <input type="text" value="Symptom"/> Aggregation: <input type="text" value="Diagnosis"/>

+ Add Feature

Clear Save

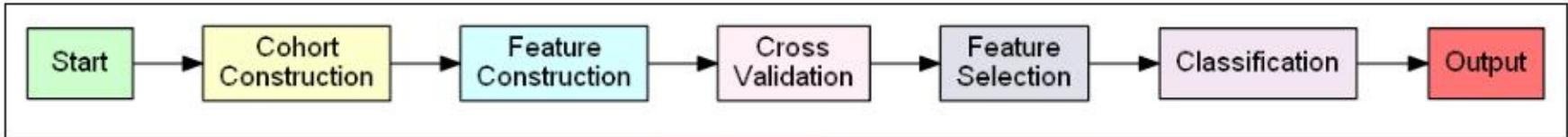
- Diagnosis
- Medication
- Lab
- Procedure
- Symptom**

PARAMO

▼ Predictive Modeling Pipeline

Pipeline: Example 1 Actions

Configuration Graph Status/Results



Cohort Construction Feature Construction Cross Validation Feature Selection Classification Other

Parameter	Value
Feature Selection	Method: Information Gain
	Number of Diagnosis: 150
	Number of Lab: 100
	Number of Medication: 50
	Number of Symptom: 20
Feature Selection	Method: Fisher Score
	Number of Diagnosis: 150
	Number of Lab: 100
	Number of Medication: 50
	Number of Symptom: 20

+ Add Feature Selection

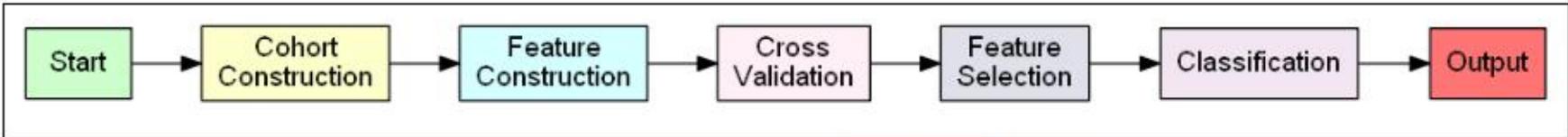
Clear Save

PARAMO

▼ Predictive Modeling Pipeline

Pipeline: Example 1 Actions

Configuration Graph Status/Results



Cohort Construction Feature Construction Cross Validation Feature Selection Classification Other

Parameter	Value	
Mode	Evaluation	
Classifier	Algorithm	Random Forest
	Parameters:	100
Classifier	Algorithm	Logistic Regression
	Parameters:	
Classifier	Algorithm	Naive Baysian
	Parameters:	Random Forest
+ Add Classifier		

Clear Save

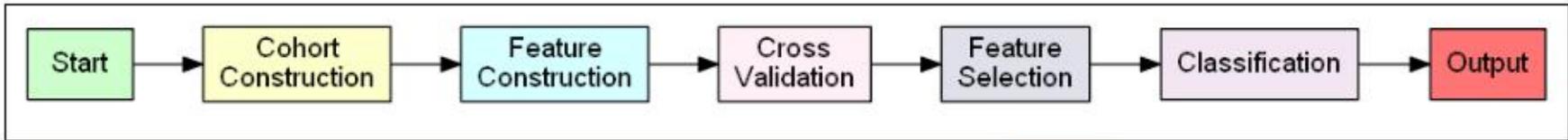
- Random Forest
- Logistic Regression
- KNN
- SVM
- Naive Baysian**

PARAMO

▼ Predictive Modeling Pipeline

Pipeline: Example 1 Actions

Configuration Graph Status/Results



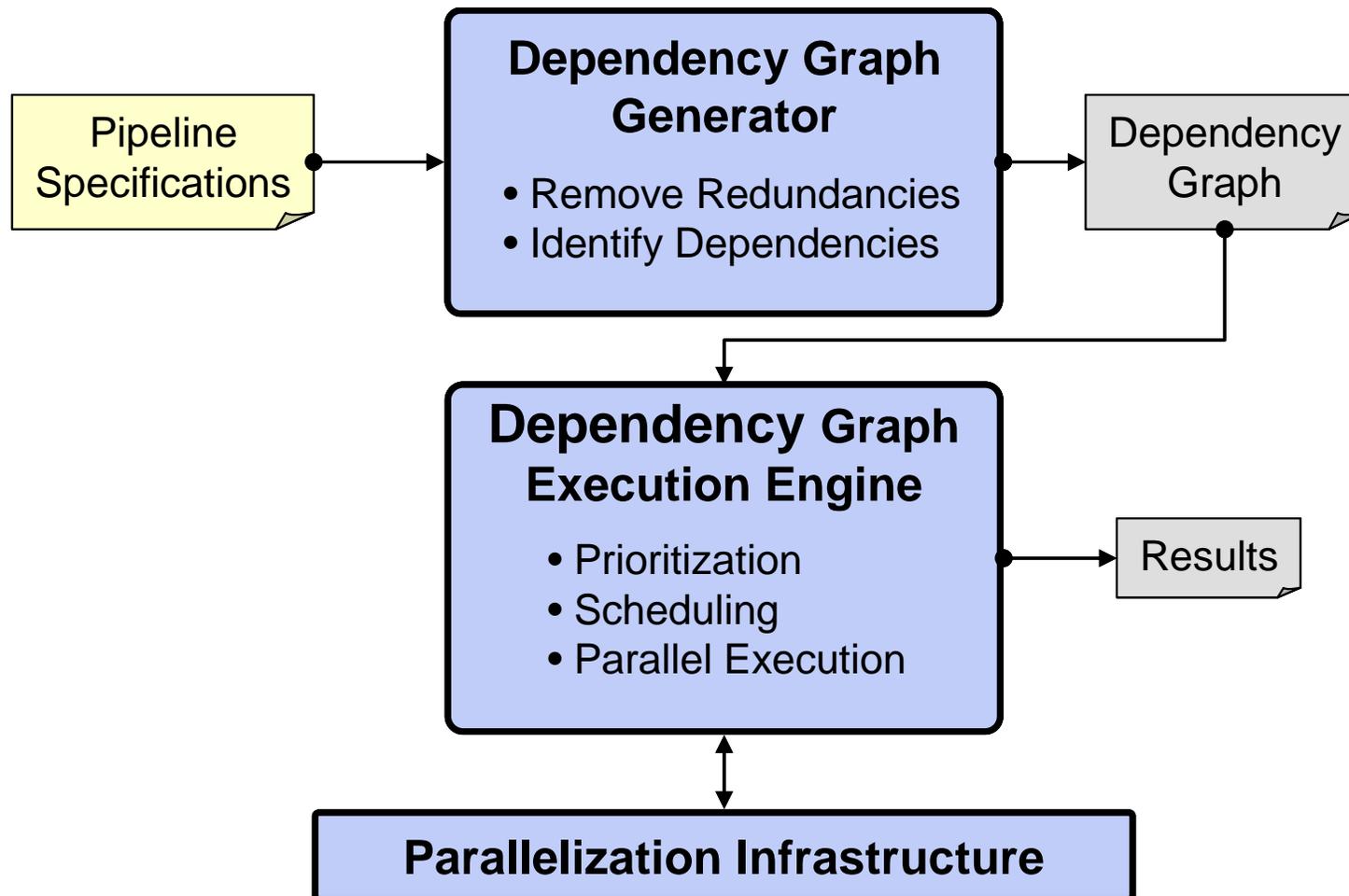
Cohort Construction Feature Construction Cross Validation Feature Selection Classification Other

Parameter	Value
Number of Parallel Jobs	20
Priority	Time

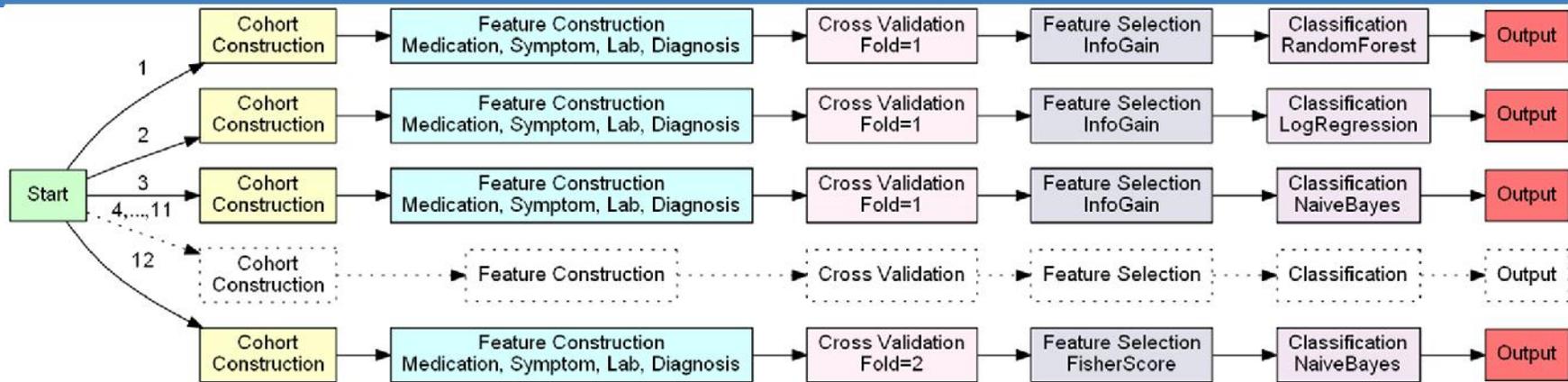
Clear Save

- None
- Time**
- Accuracy

PARAMO: A Parallel Predictive Modeling Platform



An Example Set of Pipeline Specifications



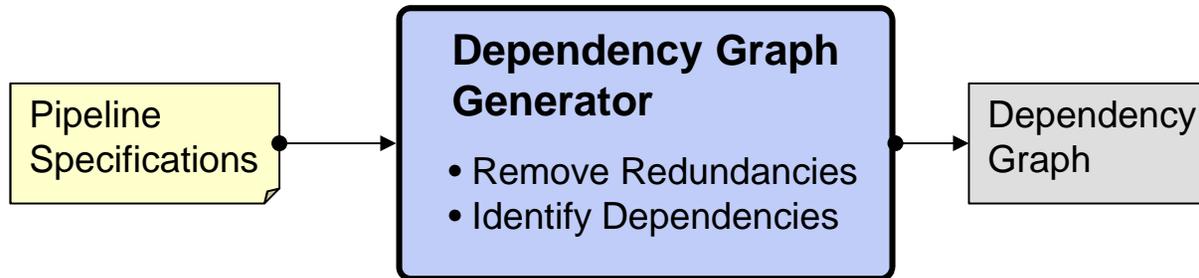
- Cohort construction: One patient data set

- Feature construction:

Feature Type	Aggregation
Diagnoses	Count
Medications	Count
Symptoms	Count
Labs	Mean

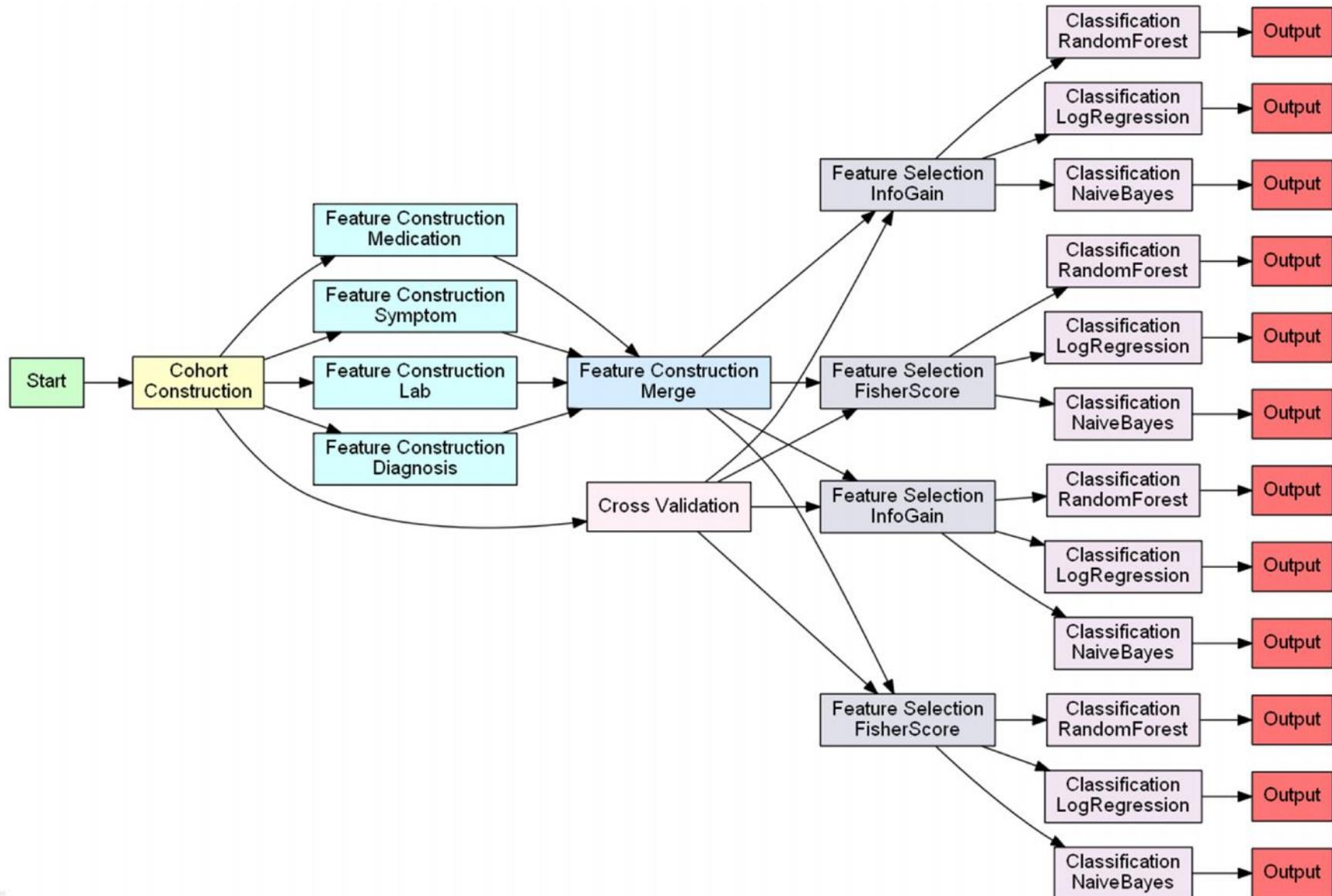
- Cross-validation: 2-fold cross-validation
- Feature selection: Information Gain, Fisher Score
- Classification: Naïve Bayes, Logistic Regression, Random Forest

Dependency Graph Generator

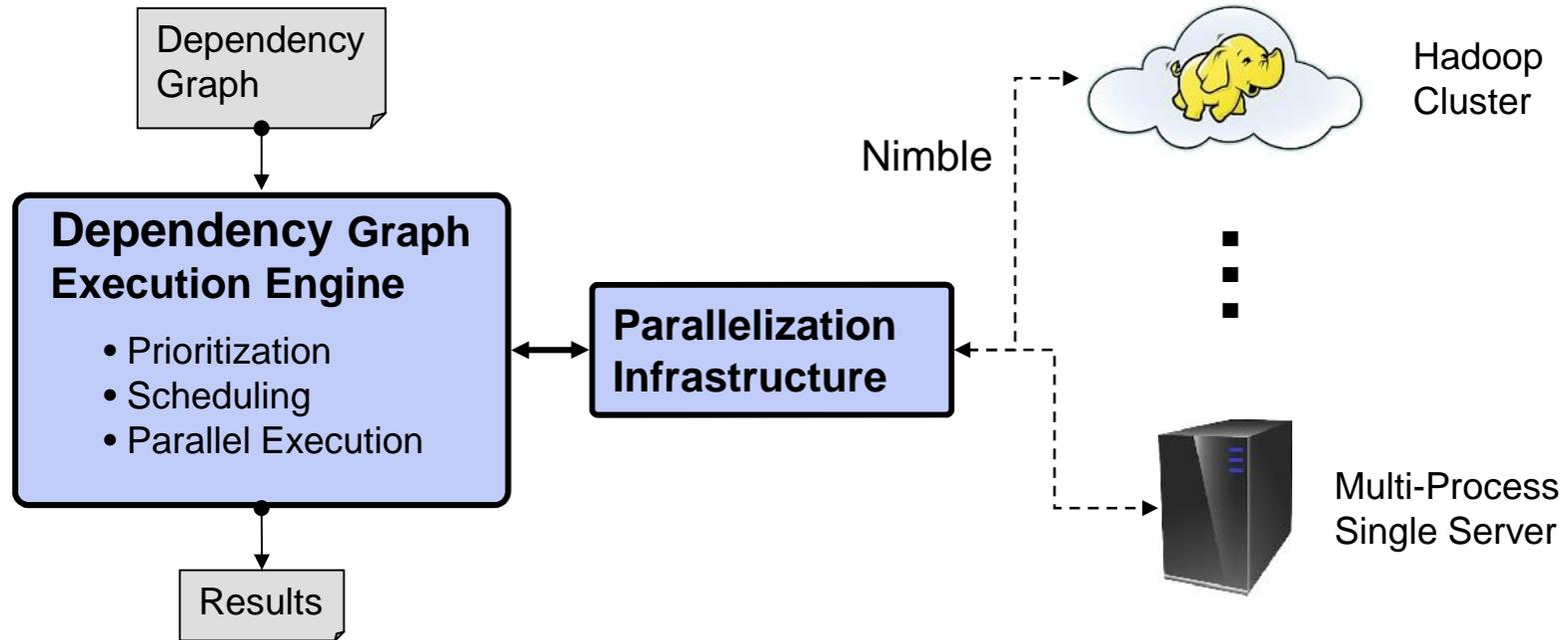


- Input: Pipeline specifications
- Output: Dependency graph
- Function:
 - Remove Redundancies
 - Identify Dependencies
 - Encode parallel jobs

Dependency Graph

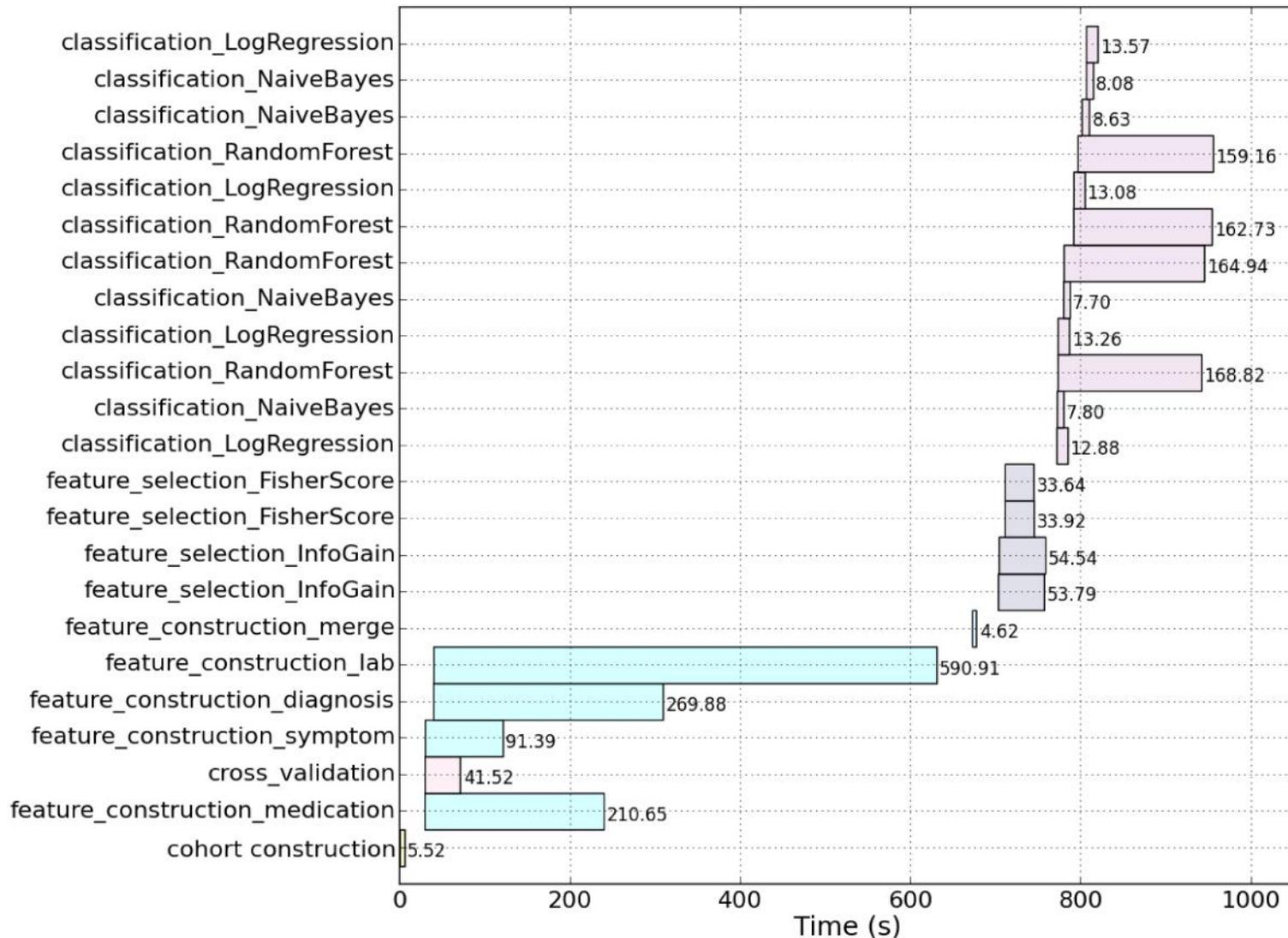


Dependency Graph Execution Engine



- Input: Dependency graph
- Output: Results (models, scores, etc.)
- Function:
 - Schedules tasks in a topological ordering of the graph
 - Prioritizes pending tasks using information from already completed tasks
 - Executes tasks in parallel via the parallelization infrastructure

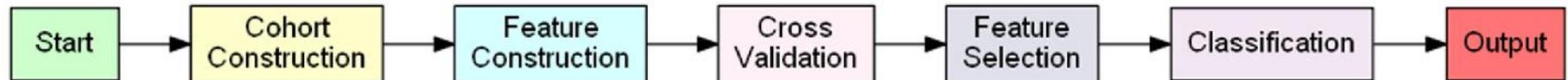
Dependency Graph Runtime Analysis (Hadoop, 20 Concurrent Tasks)



Experimental Data Sets

Data Set	Years of Data	Number of Patients	Number of Features	Number of Records	Number of Cases	Number of Controls	Target Condition
Small	3	4,758	25932	3,312,558	615	949	Hypertension Control
Medium	10	32,675	46117	24,719,809	4644	28031	Heart Failure Onset
Large	4	319,650	49269	33,531,311	16385	164743	Hypertension Onset

Experimental Pipeline Specifications



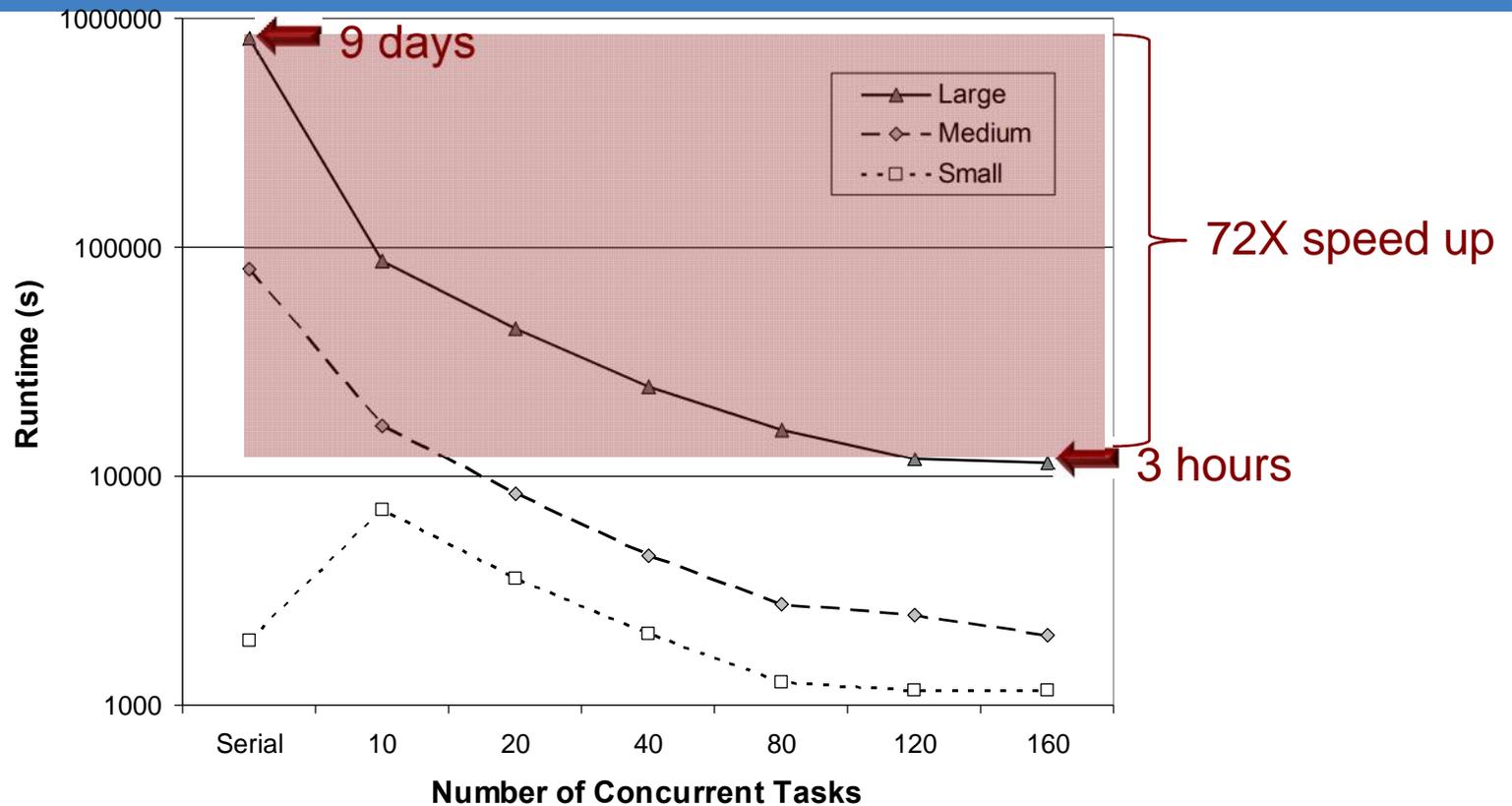
- Cohort construction: One patient data set: Small, Medium, or Large

- Feature construction:

Feature Type	Aggregation
Diagnoses	Count
Medications	Count
Procedures	Count
Symptoms	Count
Labs	Mean

- Cross-validation: 10 x 10-fold cross-validation
- Feature selection: Information Gain, Fisher Score
- Classification: k-NN, Naïve Bayes, Logistic Regression, Random Forest

Running Time vs. Parallelism level



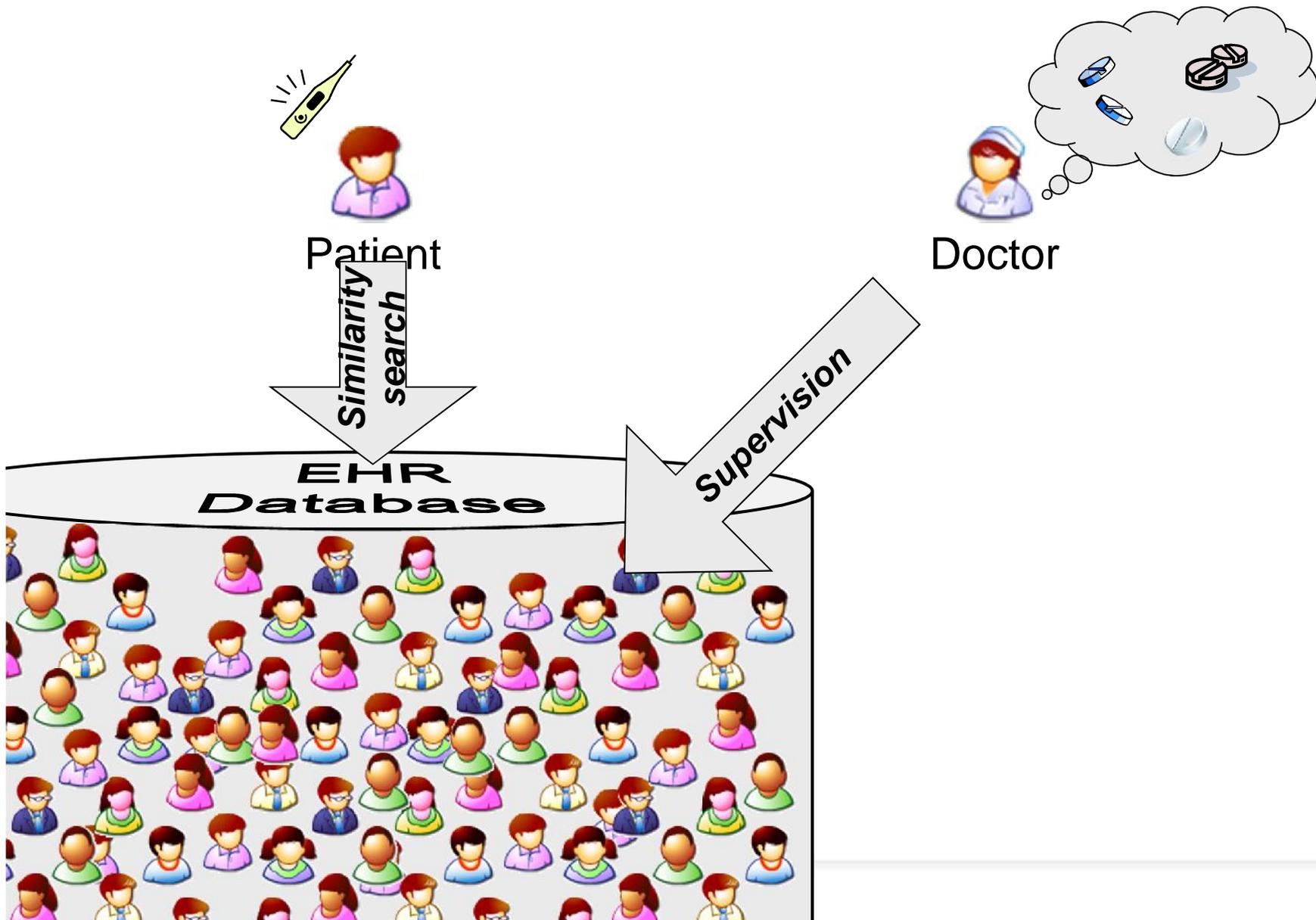
- Small: 5,000 patients, Medium: 33K, Large: 319K
- 10 times 10-fold cross validation
- Dependency graph: 1808 nodes and 3610 edges

Summary

- Predictive models in healthcare research is becoming more prevalent
- Electronic health records (EHR) adoption continues to accelerate
- Need for scalable predictive modeling platforms/systems
- PARAMO is a parallel predictive modeling platform for EHR data
- PARAMO can facilitate large-scale modeling endeavors and speed-up the research workflow
- Tests on real EHR data show significant performance gains

PATIENT SIMILARITY PLATFORM

Patient Similarity Problem



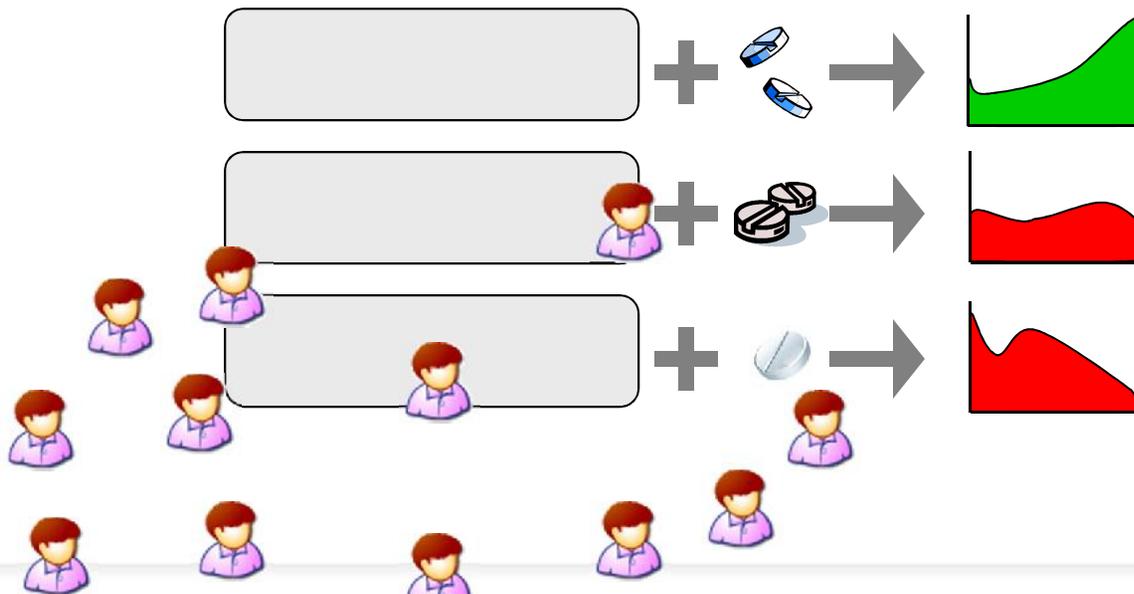
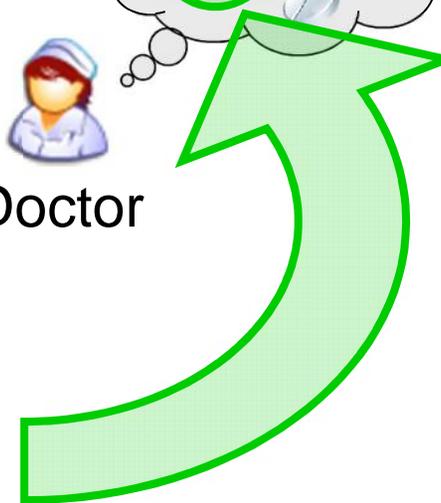
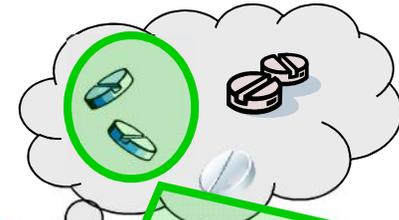
Patient Similarity Problem



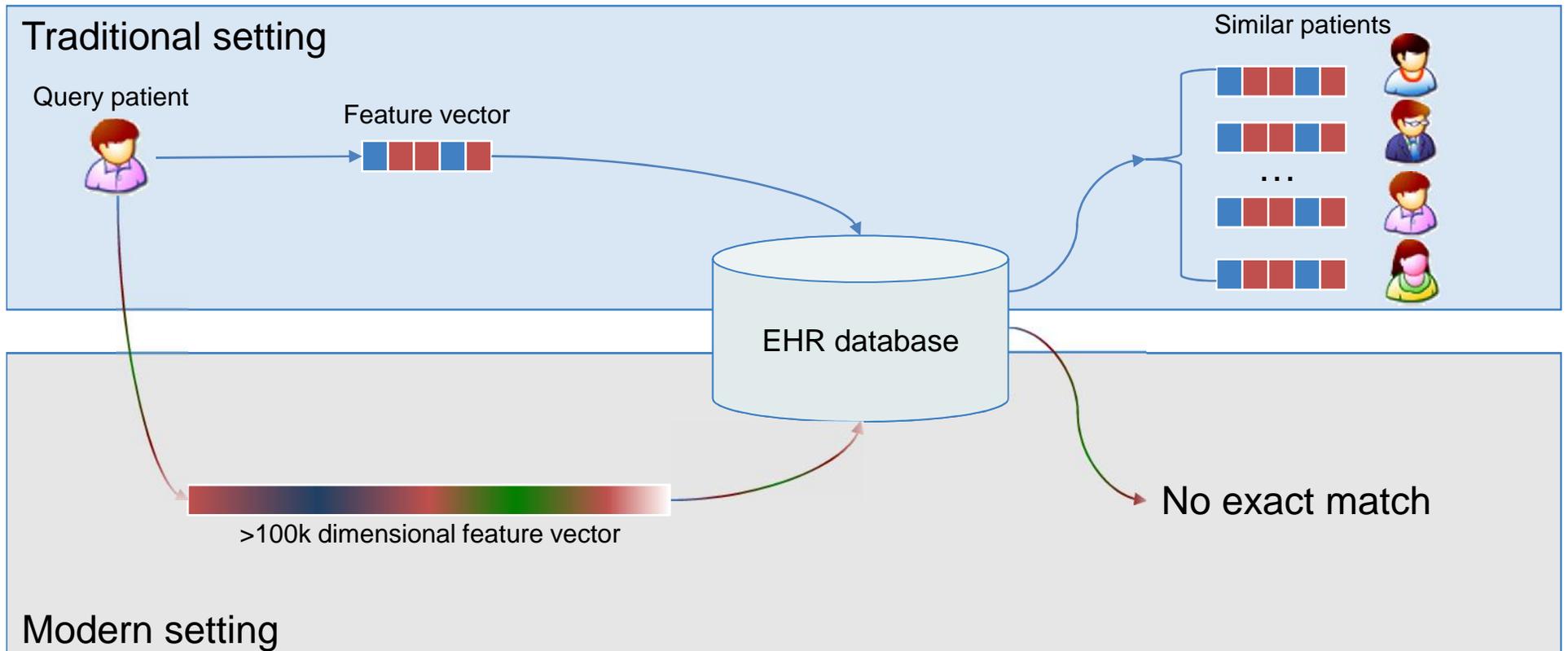
Patient



Doctor

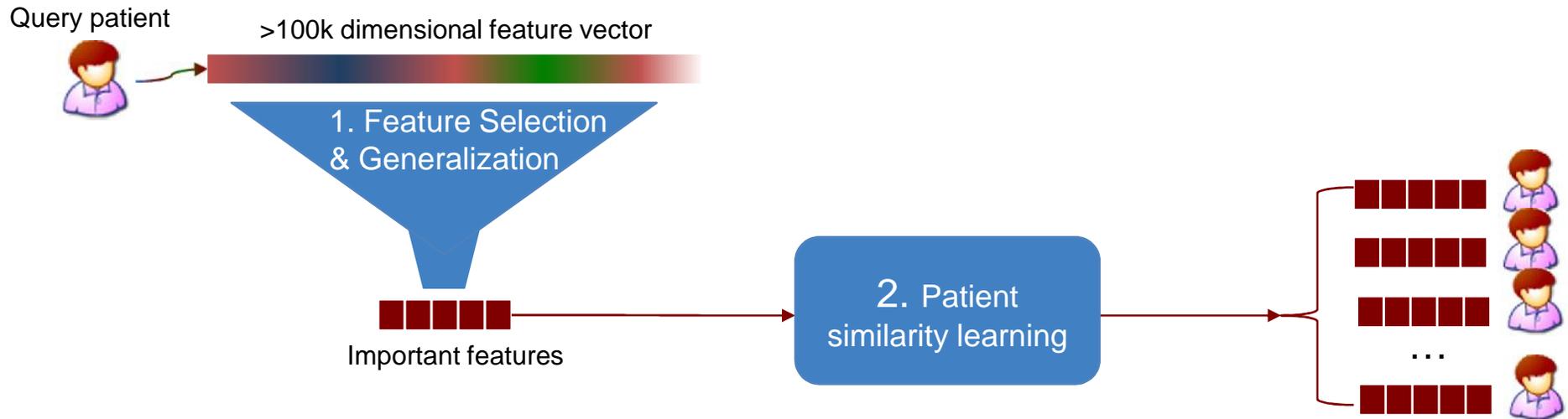


Challenges of Patient Similarity



- As the size of a feature vector increases, it is hard to find exact match on all features
- How to find relevant patients to a query for a specific clinical context?

Our Approach



- For a clinical context,
 1. What are important features?
 2. What is the right similarity measure?

Large-scale Analytics Platform

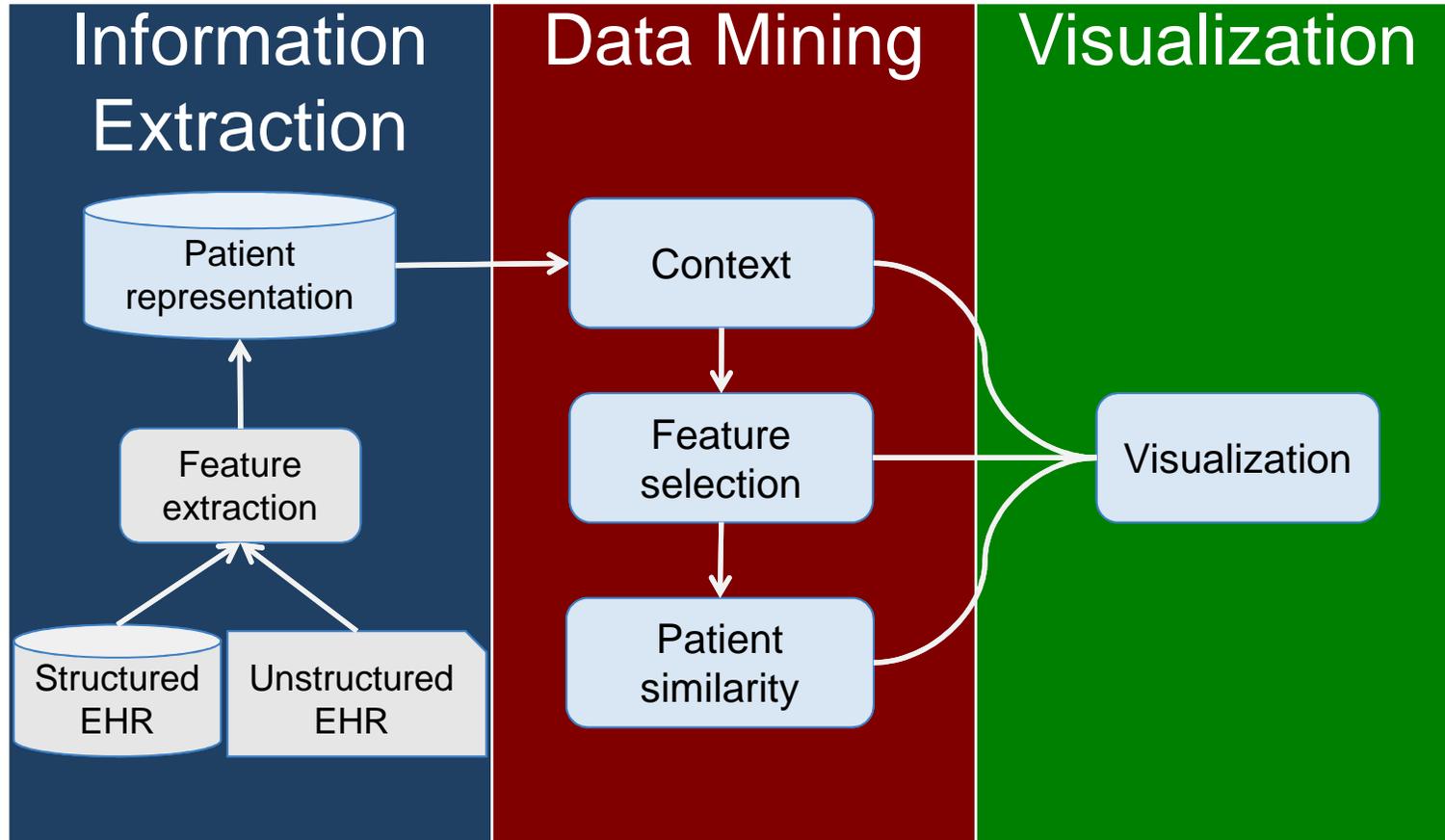
Healthcare Analytic Platform

Information
Extraction

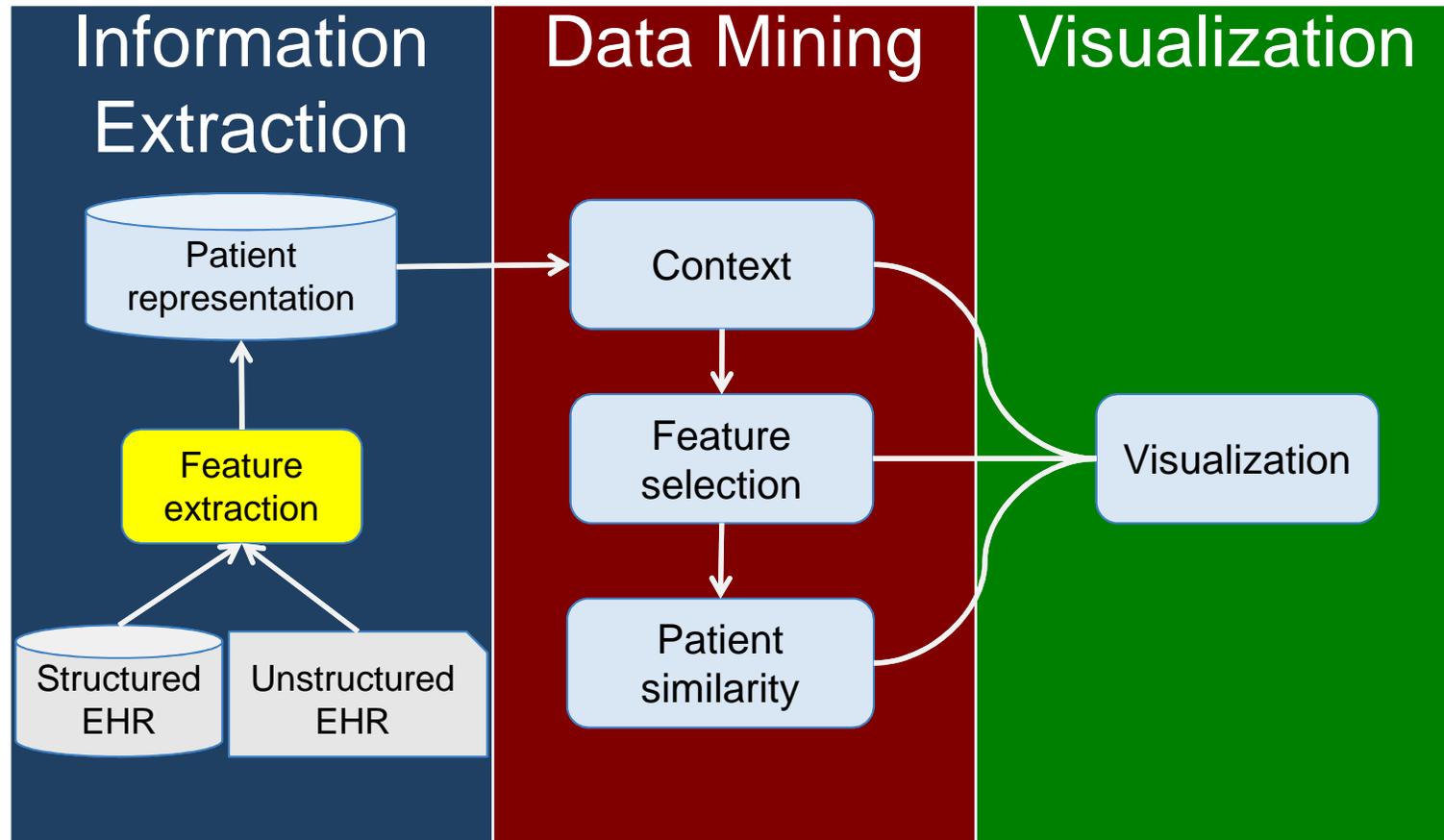
Data Mining

Visualization

Healthcare Analytic Platform



Feature Extraction from Unstructured EHR data



Motivations for Early Detection of Heart Failure

- Heart failure (HF) is a complex disease
- Huge Societal Burden

5 millions

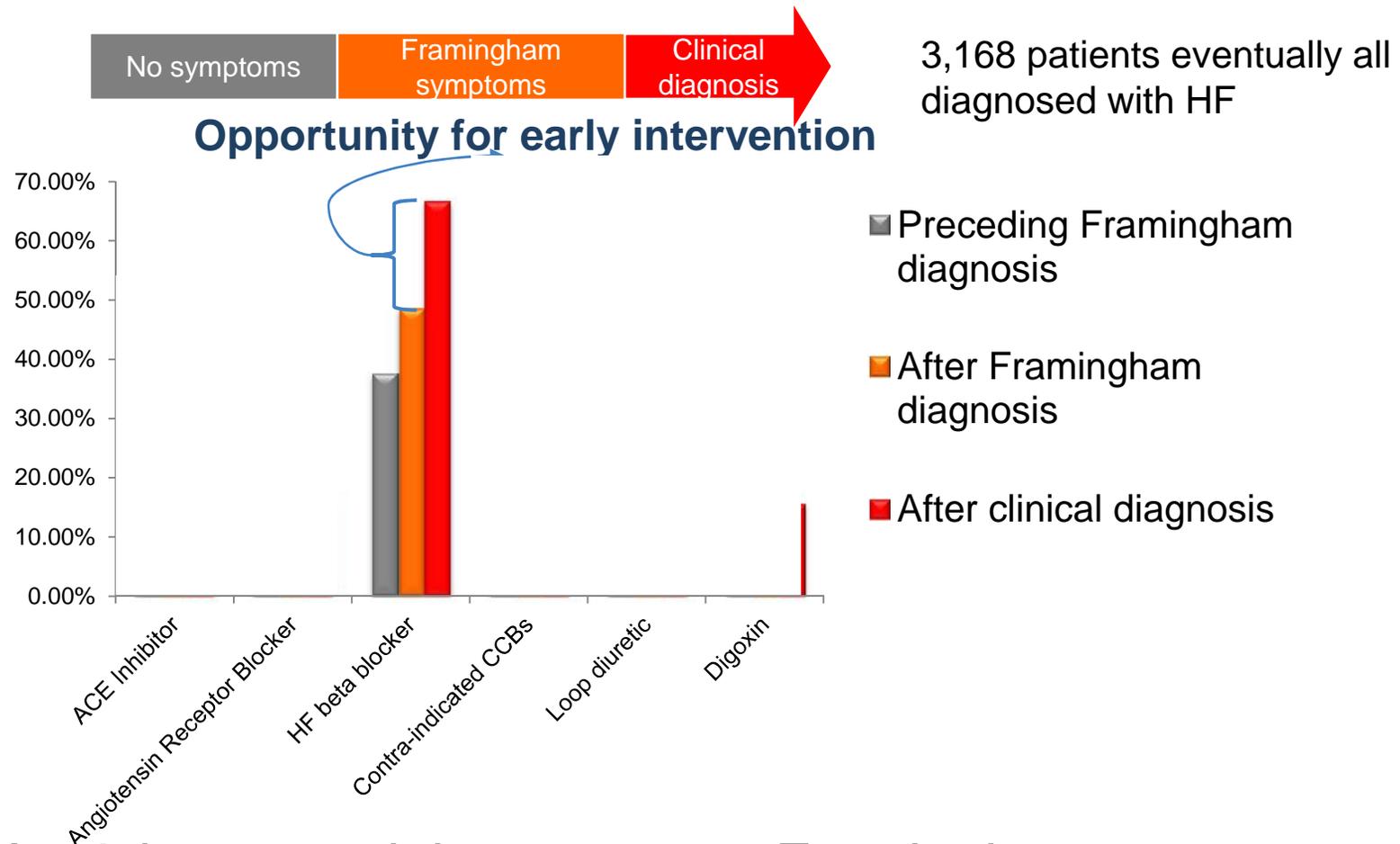
0.5 millions

20%

48%

- Diagnoses are usually made late, despite there are symptoms documented in clinical notes prior
- Our method extracting HF symptoms achieves precision 0.925, recall 0.896, and *F*-score 0.910

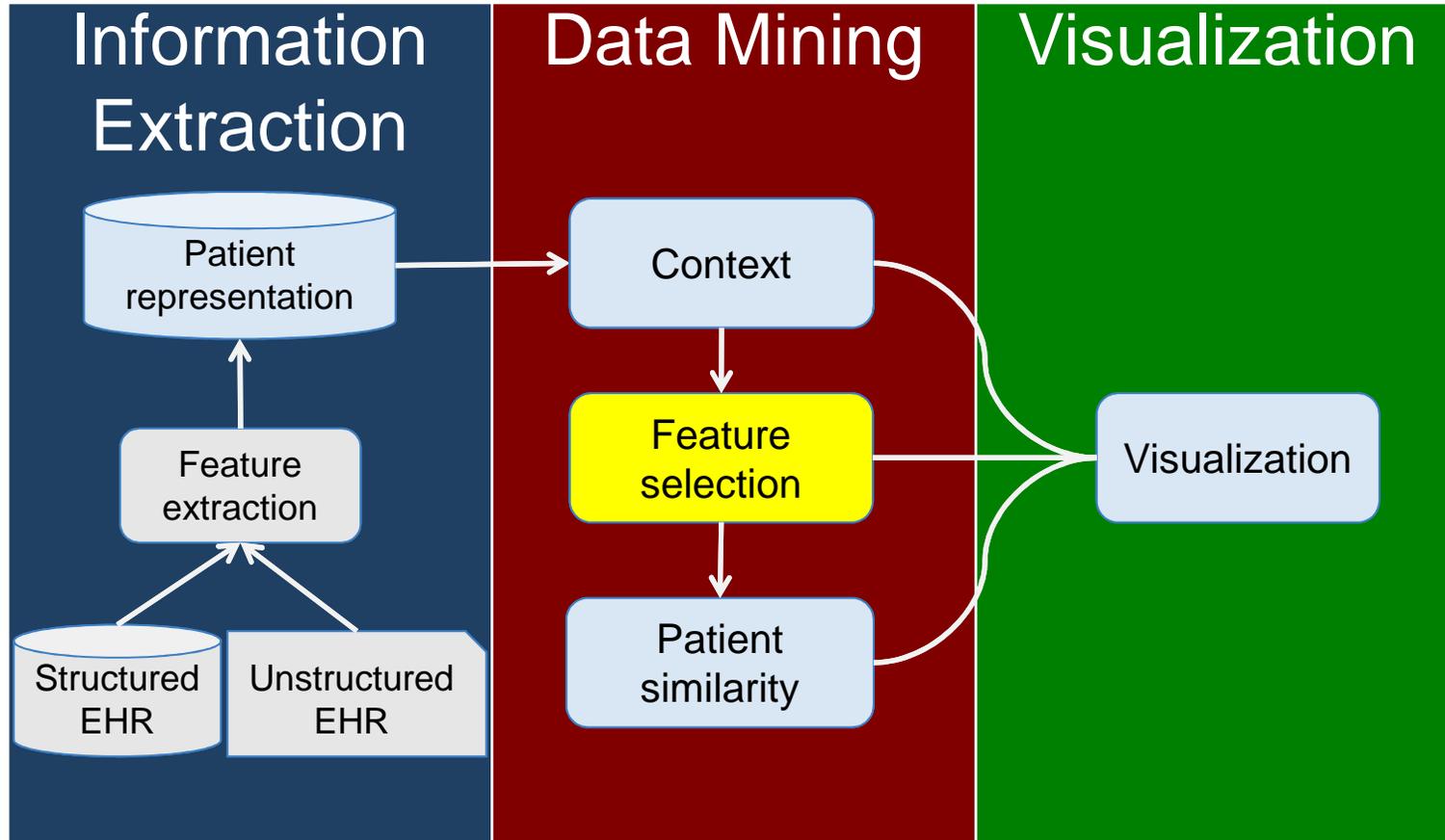
Potential Impact on Evidence-based Therapies



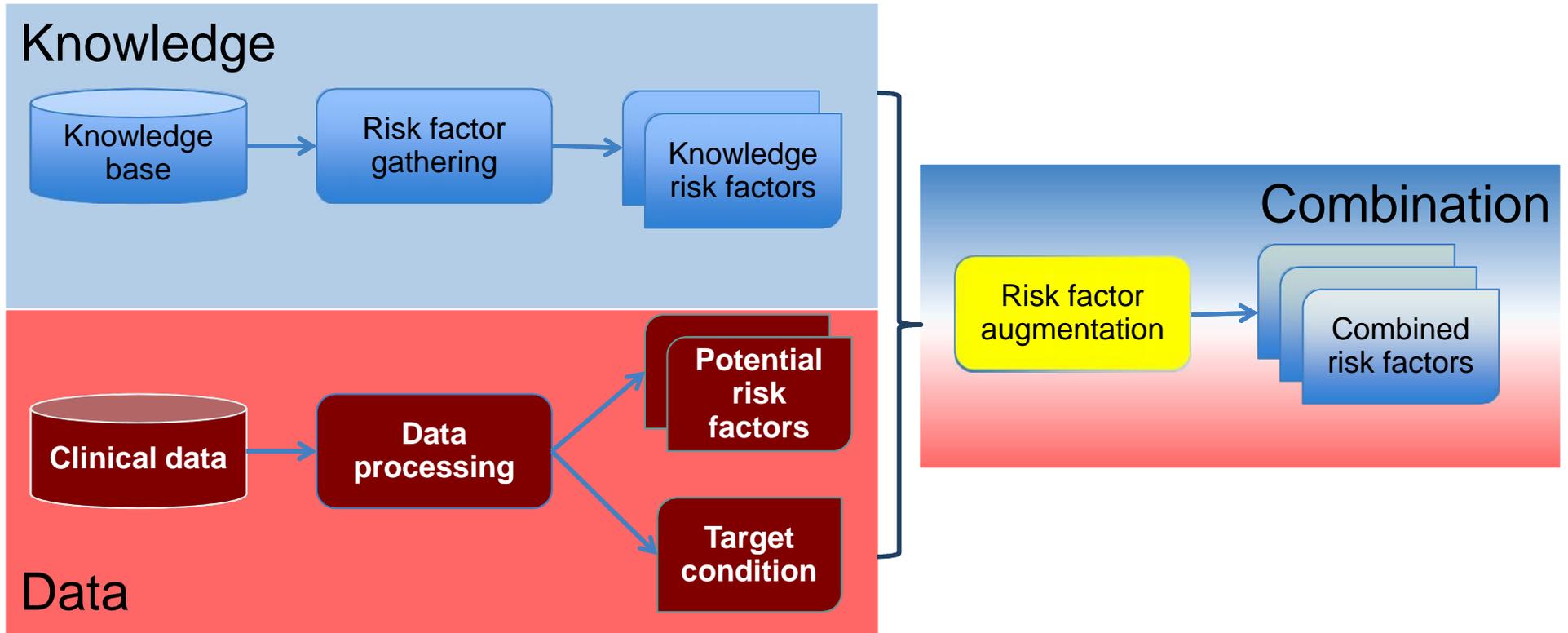
- Applying text mining to extract Framingham symptoms can help trigger early intervention

Vhavakrishnan R, Steinhubl SR, Sun J, et al. Potential impact of predictive models for early detection of heart failure on the initiation of evidence-based therapies. J Am Coll Cardiol. 2012;59(13s1):E949-E949.

Knowledge plus Data Feature Selection



Combining Knowledge- and Data-driven Risk Factors



Jimeng Sun, Jianying Hu, Dijun Luo, Marianthi Markatou, Fei Wang, Shahram Ebadollahi, Steven E. Steinhubl, Zahra Daar, Walter F. Stewart. Combining Knowledge and Data Driven Insights for Identifying Risk Factors using Electronic Health Records. AMIA2012

Dijun Luo, Fei Wang, Jimeng Sun, Marianthi Markatou, Jianying Hu, Shahram Ebadollahi, SOR: Scalable Orthogonal Regression for Low-Redundancy Feature Selection and its Healthcare Applications. SDM'12

Risk Factor Augmentation

- Sparse learning objective formulation:

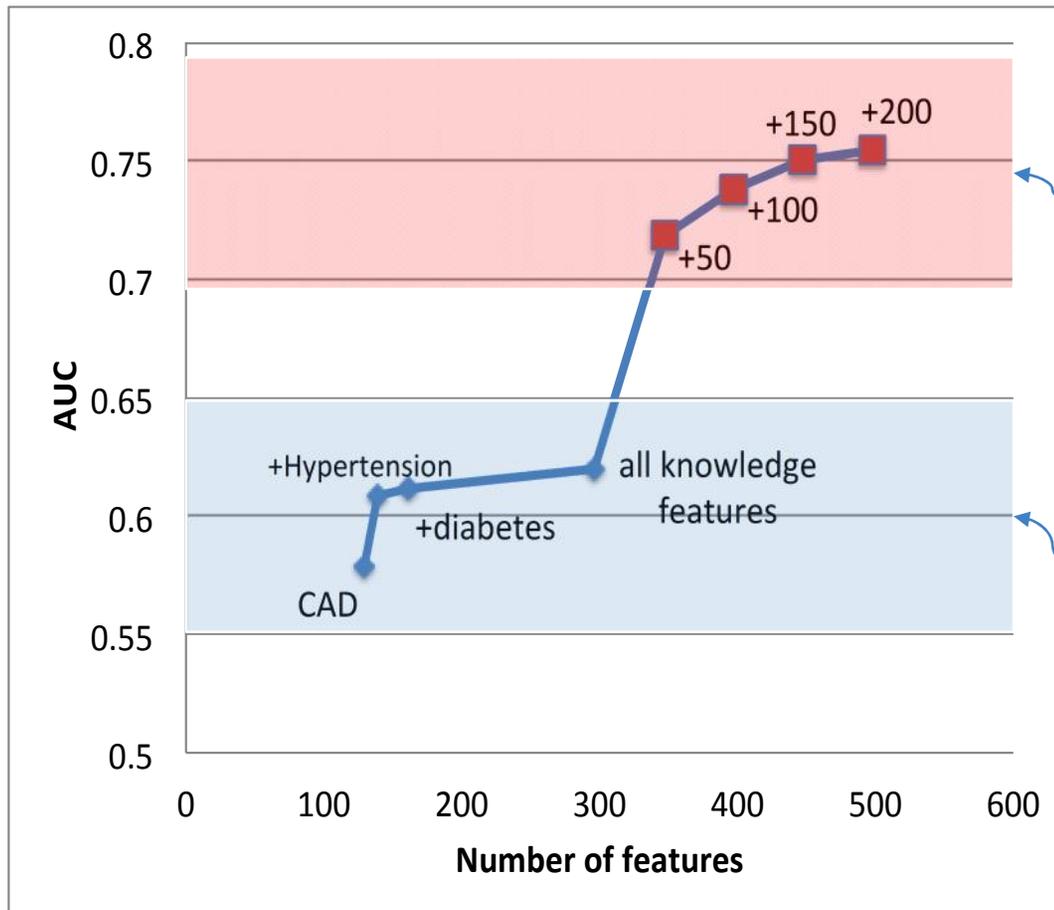
$$f(\boldsymbol{\alpha}) = \frac{1}{2} \underbrace{\|\mathbf{y} - \mathbf{X}\boldsymbol{\alpha}\|^2}_{\text{Model error}}$$

Jimeng Sun, Jianying Hu, Dijun Luo, Marianthi Markatou, Fei Wang, Shahram Ebadollahi, Steven E. Steinhubl, Zahra Daar, Walter F. Stewart. Combining Knowledge and Data Driven Insights for Identifying Risk Factors using Electronic Health Records.

AMIA2012

Dijun Luo, Fei Wang, Jimeng Sun, Marianthi Markatou, Jianying Hu, Shahram Ebadollahi, SOR:
Scalable Orthogonal Regression for Low-Redundancy Feature Selection and its Healthcare Applications. SDM'12

Prediction Results using Selected Features



Data driven features improve AUC significantly

Knowledge driven features improve AUC slightly

Top-10 Selected Data-driven Features

Feature	Relevancy to HF
Dyslipidemia	
Thiazides-like Diuretics	
Antihypertensive Combinations	
Aminopenicillins	
Bone density regulators	
Natrietic Peptide	
Rales	
Diuretic Combinations	
S3Gallop	
NSAIDS	

- 9 out of 10 are considered relevant to HF

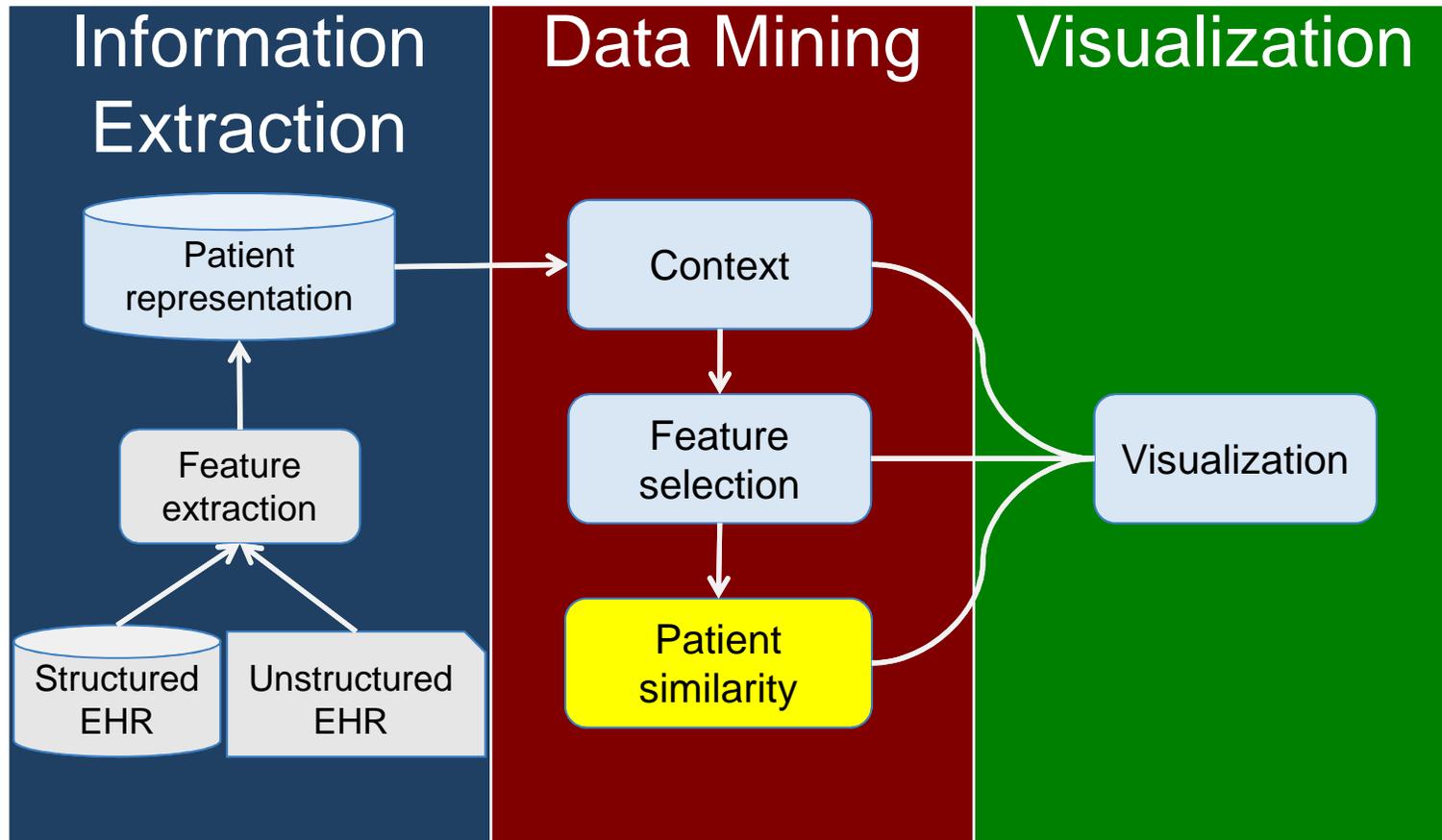
Top-10 Selected Data-driven Features

Category
Diagnosis
Medication
Lab
Symptom

Feature	Relevancy to HF
Dyslipidemia	
Thiazides-like Diuretics	
Antihypertensive Combinations	
Aminopenicillins	
Bone density regulators	
Natrietic Peptide	
Rales	
Diuretic Combinations	
S3Gallop	
NSAIDS	

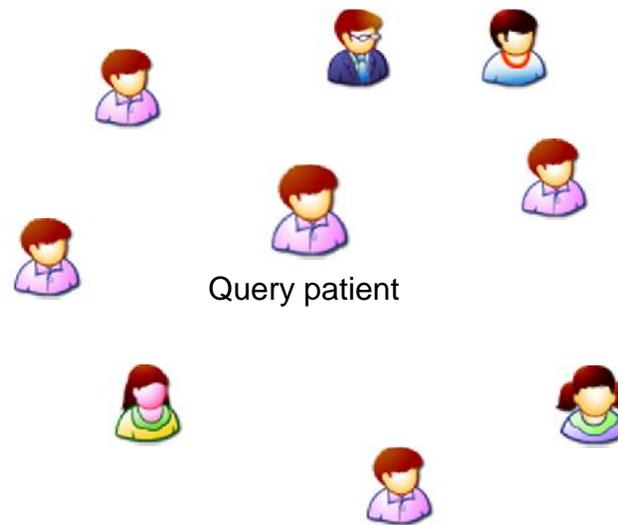
- 9 out of 10 are considered relevant to HF
- The data driven features are complementary to the existing knowledge-driven features

Patient Similarity



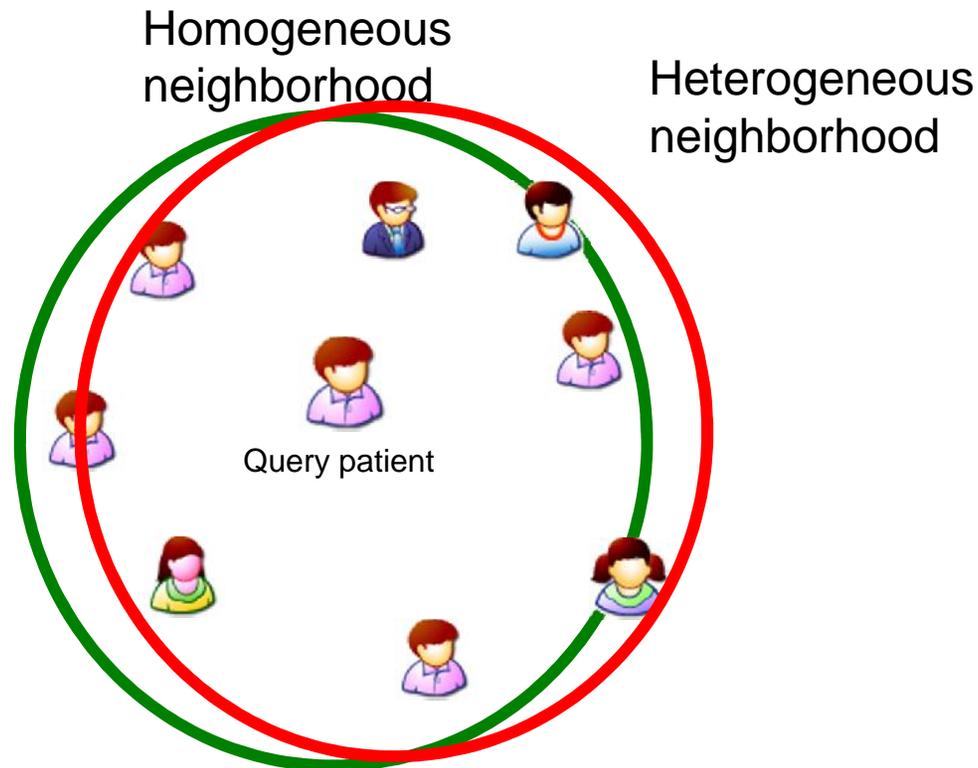
Patient Similarity through Locally Supervised Metric Learning

Under a specific clinical context



Patient Similarity through Locally Supervised Metric Learning

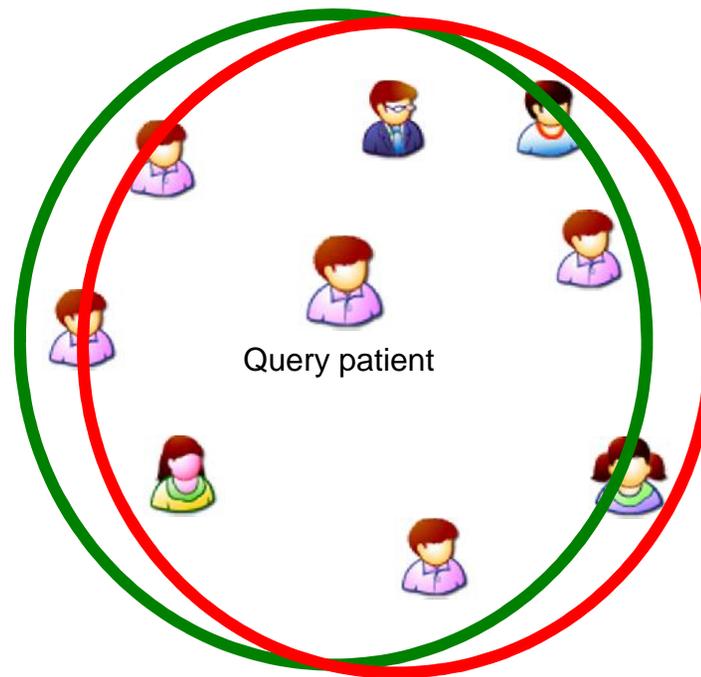
Under a specific clinical context



- Homogeneous neighbors: true positives
- Heterogeneous neighbors: false positives

Patient Similarity through Locally Supervised Metric Learning

Under a specific clinical context



- Shrink homogeneous neighborhood
- Grow heterogeneous neighborhood

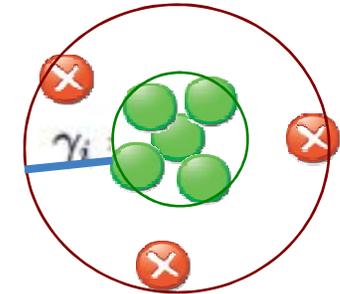
Locally Supervised Metric Learning (LSML)

Goal: Learn a generalized Mahalanobis distance for a specific clinical context (target label)

$$d_{\Sigma}(\mathbf{x}_i, \mathbf{x}_j) = \sqrt{(\mathbf{x}_i - \mathbf{x}_j)^{\top} \Sigma (\mathbf{x}_i - \mathbf{x}_j)} \quad \Sigma = \mathbf{W}\mathbf{W}^{\top}$$

Margin for x_i

$$\gamma_i = \underbrace{\sum_{k:\mathbf{x}_k \in \mathcal{N}_i^e} \|\mathbf{W}^T \mathbf{x}_i - \mathbf{W}^T \mathbf{x}_k\|^2}_{\text{Total distance to heterogeneous neighbors}} - \underbrace{\sum_{j:\mathbf{x}_j \in \mathcal{N}_i^o} \|\mathbf{W}^T \mathbf{x}_i - \mathbf{W}^T \mathbf{x}_j\|^2}_{\text{Total distance to homogeneous neighbors}}$$



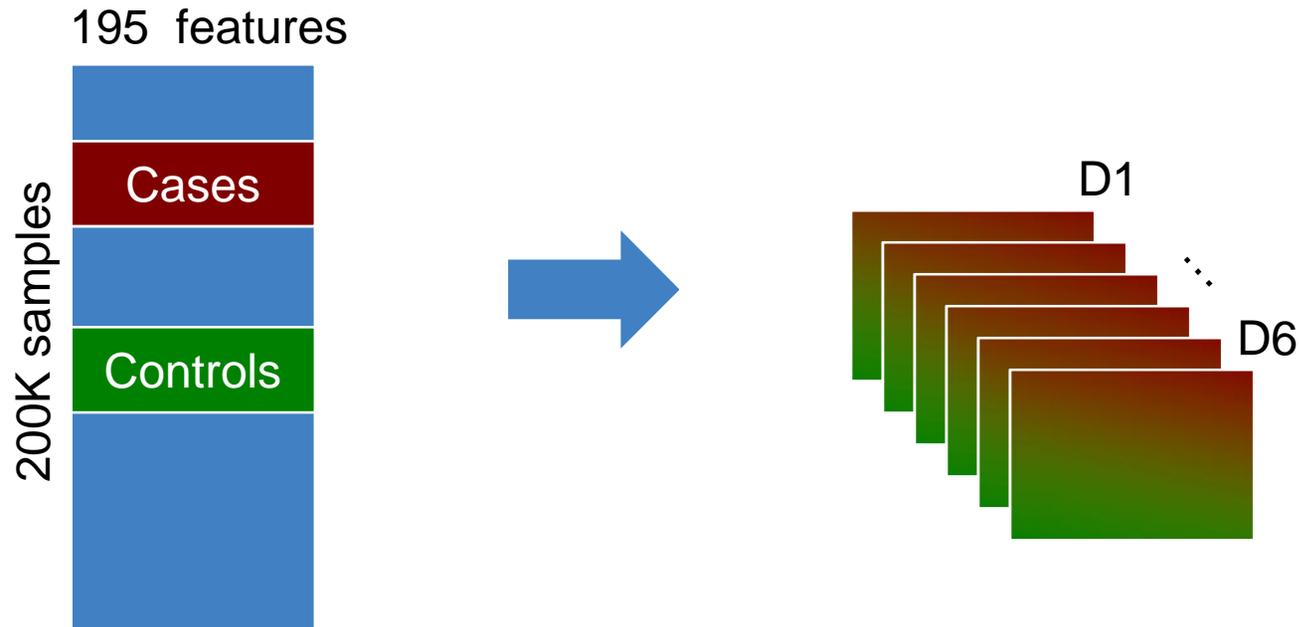
Maximize the total margin

$$\begin{aligned} \gamma = & \sum_i \sum_{k:\mathbf{x}_k \in \mathcal{N}_i^e} \mathbf{W}^T (\mathbf{x}_i - \mathbf{x}_k) (\mathbf{x}_i - \mathbf{x}_k)^T \mathbf{W} \\ & - \sum_i \sum_{j:\mathbf{x}_j \in \mathcal{N}_i^o} \mathbf{W}^T (\mathbf{x}_i - \mathbf{x}_j) (\mathbf{x}_i - \mathbf{x}_j)^T \mathbf{W} \end{aligned}$$

\mathcal{N}_i^o Homogeneous neighborhood for x_i

\mathcal{N}_i^e Heterogeneous neighborhood for x_i

Patient Similarity Experiment Design



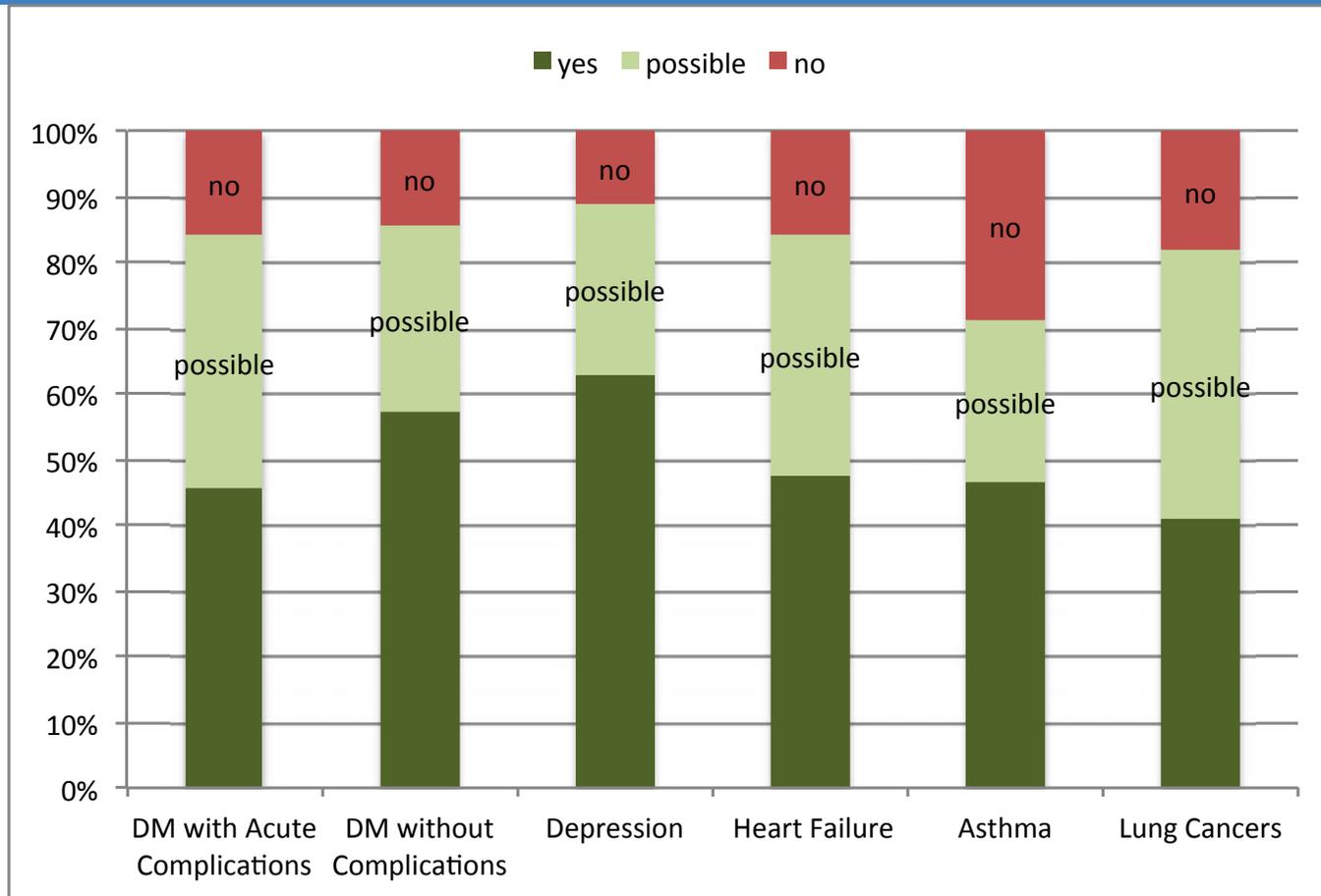
	Disease Target	samples
D1	DM with Acute Complications	4,392
D2	DM without Complications	10,734
D3	Depression	6,794
D4	Heart Failure	5,262
D5	Asthma	6,606
D6	Lung Cancers	1,172

Prediction Results on Patient Similarity

- Baselines:
 - EUC: Euclidean distance
 - PCA: Principal component analysis
 - LDA: Linear discriminant analysis
- Observations:
 - LDA does not perform well, because of the resulting dimensionality is too low
 - LSML algorithm performs the best among all

	DM with Acute Complications	DM without Complications	Depression	Congestive Heart Failure	Asthma	Lung Cancers
Euclidean	0.539	0.638	0.609	0.688	0.602	0.645
LDA	0.541	0.604	0.589	0.564	0.584	0.595
PCA	0.57	0.639	0.625	0.697	0.617	0.664
LSML	0.576	0.669	0.632	0.723	0.625	0.677

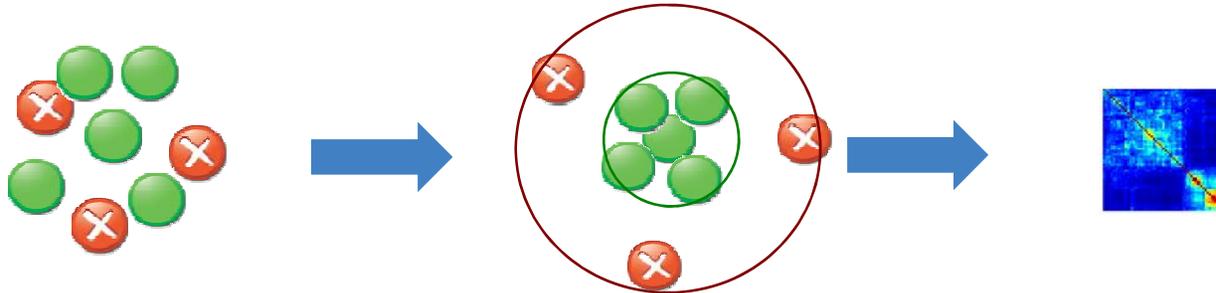
Clinical Relevancy of Patient Similarity Results



- Retrieve the top ranked comorbidities among similar patients
- **over 80% of those are considered as relevant to target disease**

Summary on Patient Similarity

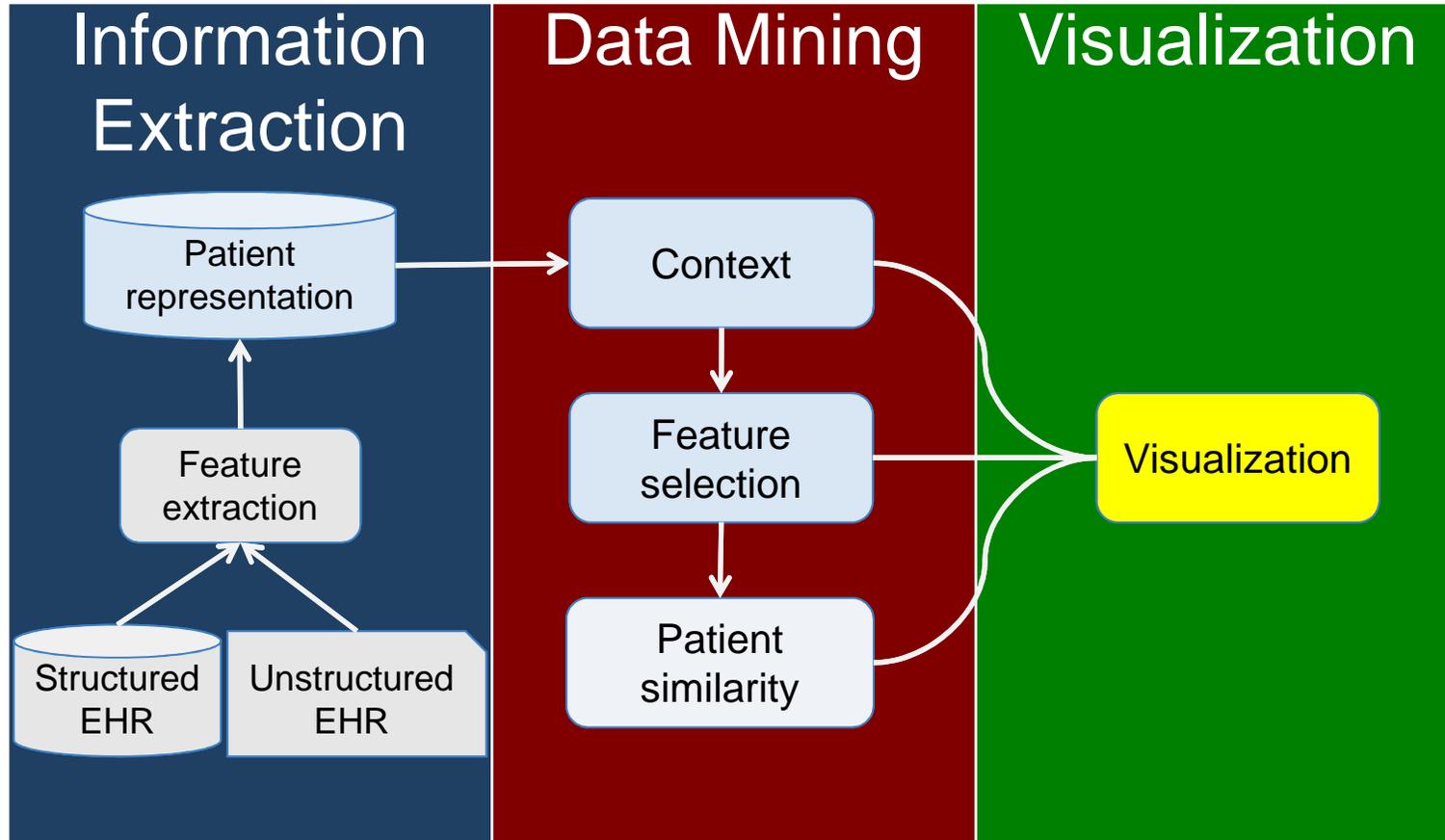
- LSML learns a customized distance metric



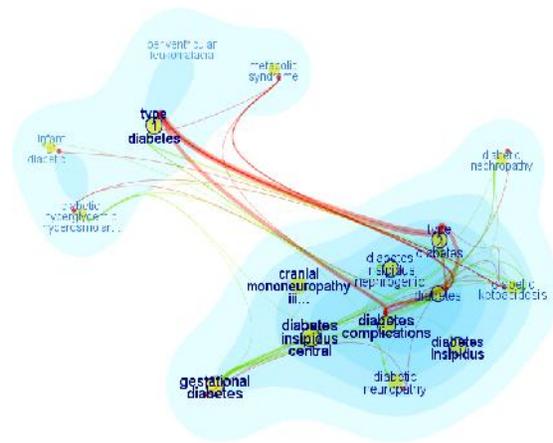
- Extension 1: Composite distance integration (Comdi) [1]
 - How to combine multiple patient similarity measures?
- Extension 2: Interactive metric update (iMet) [2]
 - How to update an existing distance measure?

1. Fei Wang, Jimeng Sun, Shahram Ebadollahi: Integrating Distance Metrics Learned from Multiple Experts and its Application in Inter-Patient Similarity Assessment. SDM 2011: 59-70 56
2. Fei Wang, Jimeng Sun, Jianying Hu, Shahram Ebadollahi: iMet: Interactive Metric Learning in Healthcare Applications. SDM 2011: 944-955

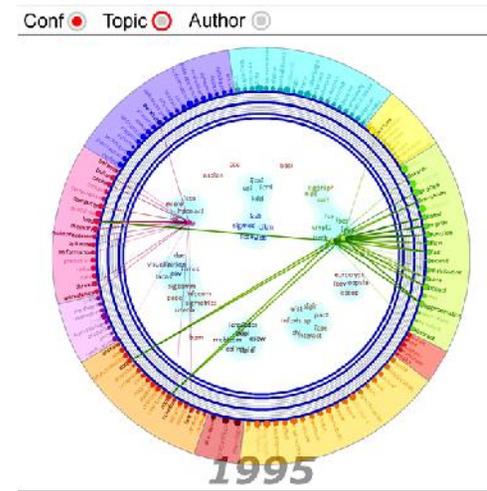
Visualization



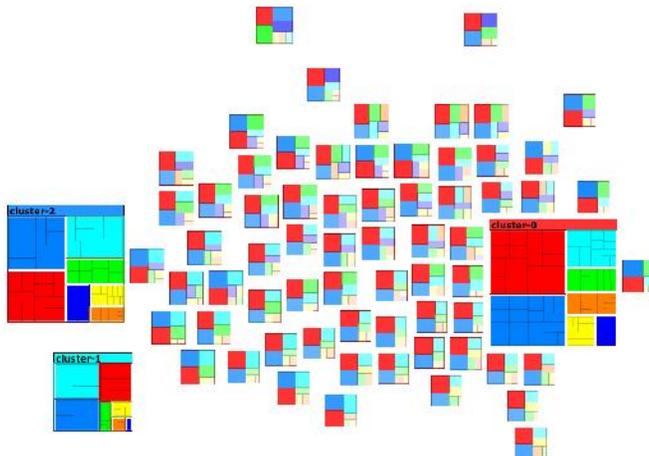
Visualization



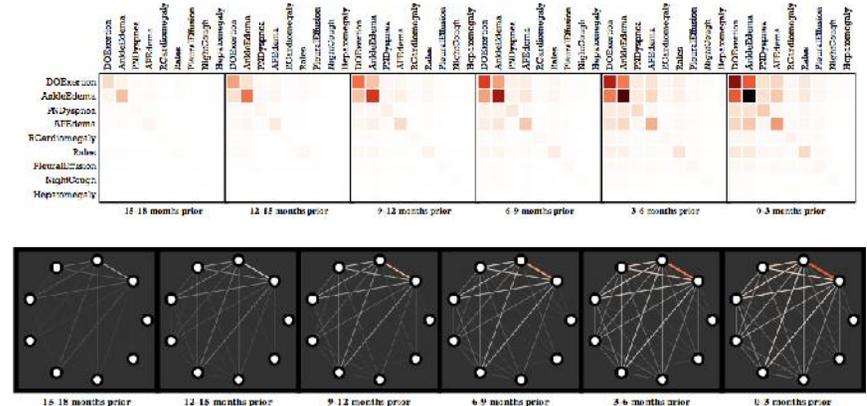
FacetAtlas (InfoVis'10)



SolarMap(ICDM'11)

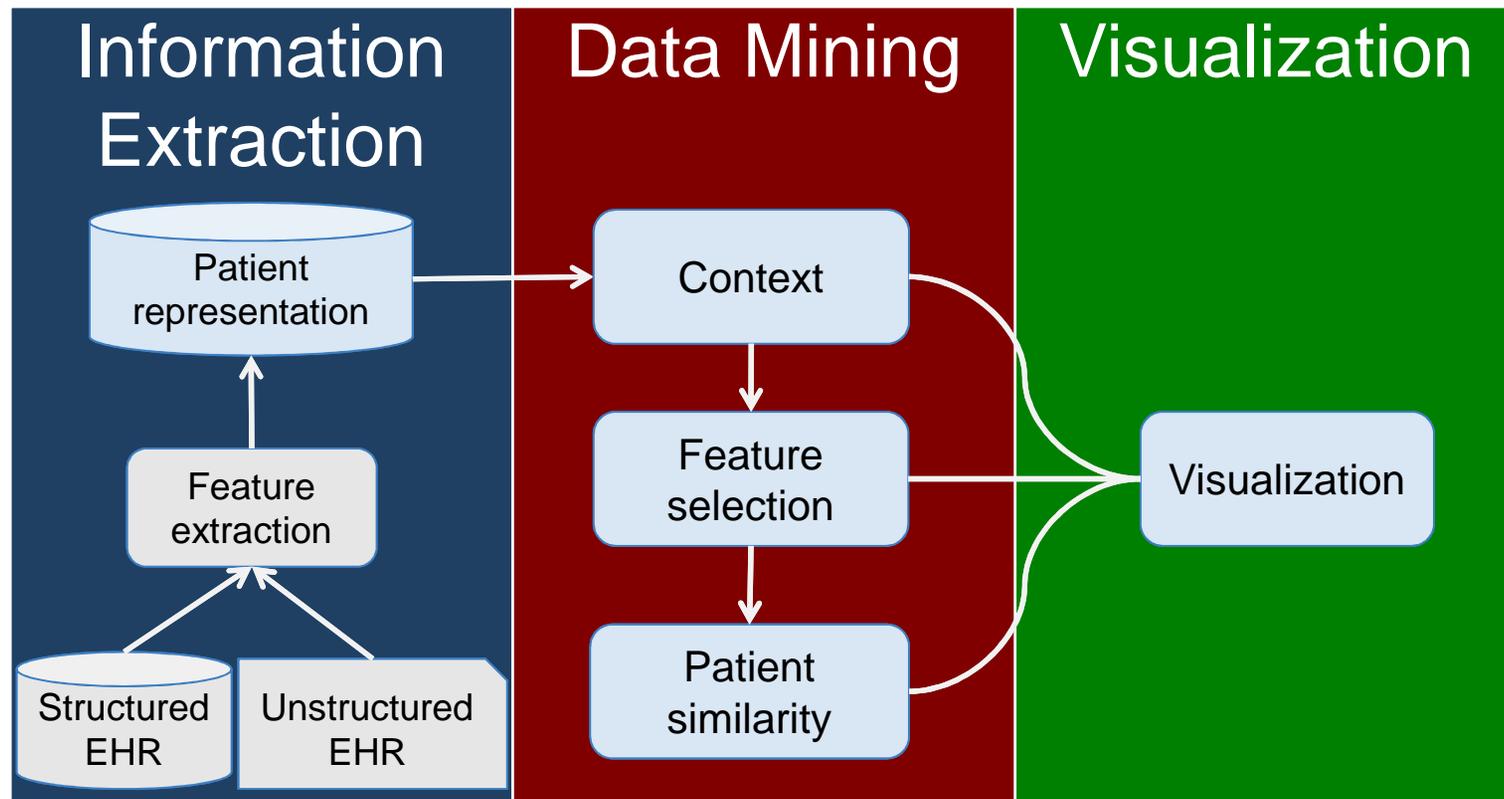


DICON (InfoVis'11)



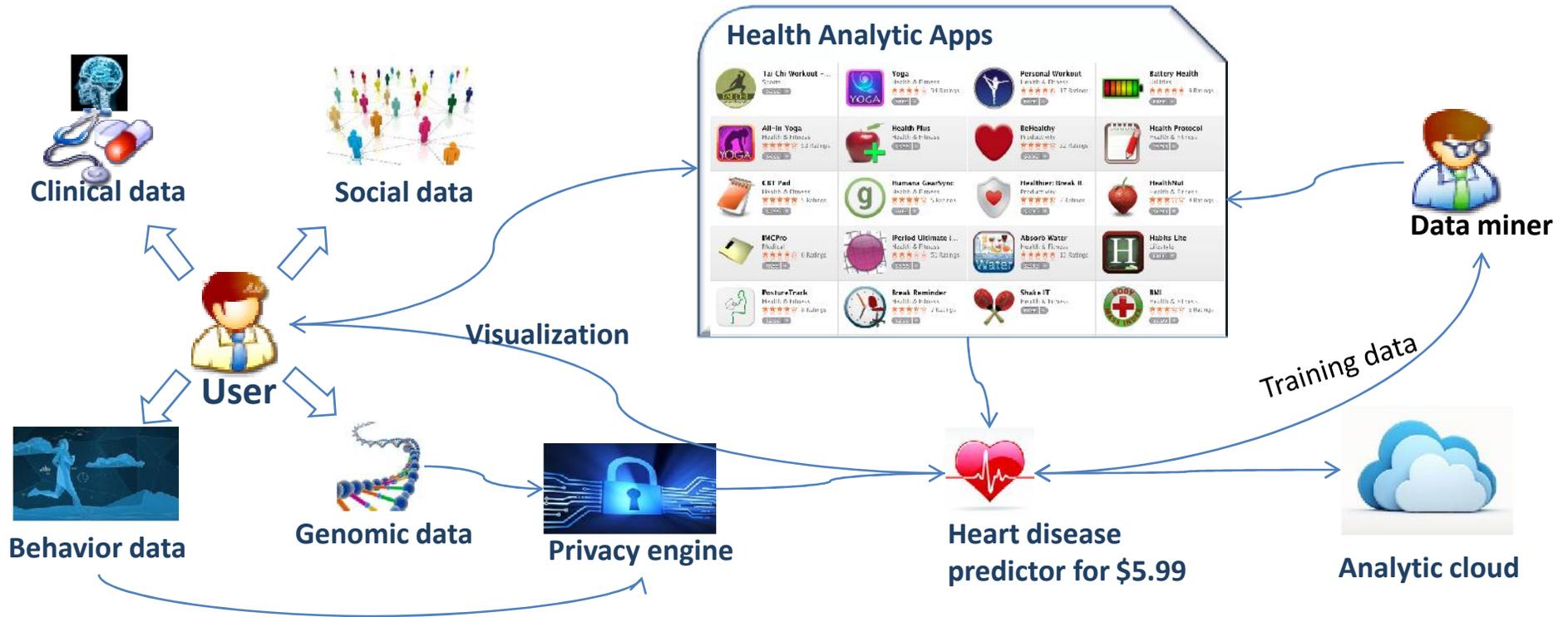
MatrixFlow (AMIA'12)

Conclusions



- Scalable healthcare analytic research platform
 - Enables efficient collaboration across domains
 - Extensible analytic platform
 - Intuitive analytic results
 - Scalable computation engine

Future of Healthcare Analytics



Research Challenges

- Data analytic techniques for each data modality
- Privacy preserving data sharing
- Visual analytic techniques