

Biomedical Science and Engineering at Oak Ridge National Laboratory

Presented by

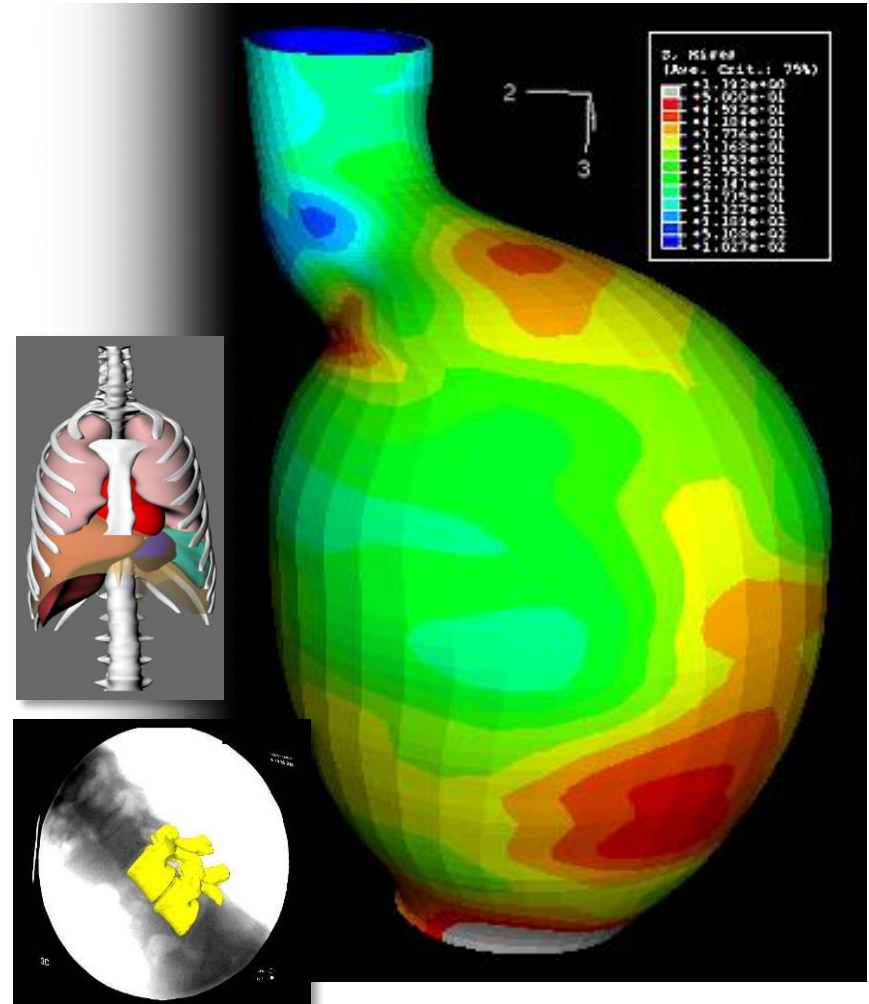
Barbara Beckerman

Modeling and Simulation Group
Computational Sciences and Engineering



Biomedical engineering and biomedical informatics research at ORNL

- Biokinetic and biotransport modeling
- Three-dimensional organ and tissue modeling using CT or other imagery (pulmonary, arterial, musculoskeletal)
- Prediction of outcomes based on biomedical models
- Knowledge discovery and intelligent agents for data mining and analysis
- Integration of models at multiple temporal and spatial scales
- Computer environments (data repositories, search tools, visualization, etc.) in support of biomedical and medical applications
- Biomedical informatics and telemedicine



Predictive multiscale modeling



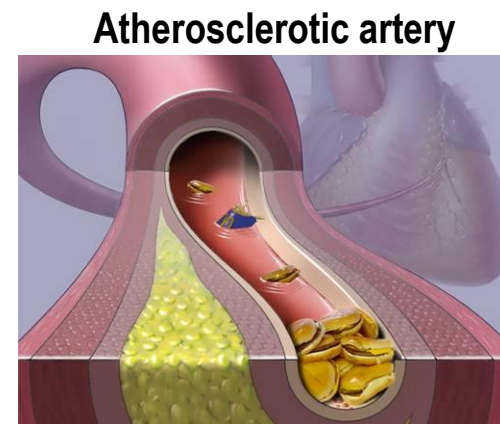
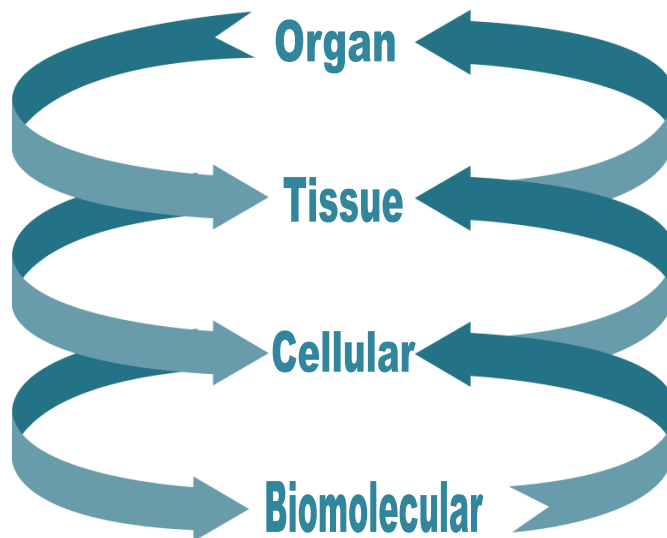
Goal: Predict migration of smooth muscle cells from media to intima due to inflammatory response after injury

Model for predicting vascular disease

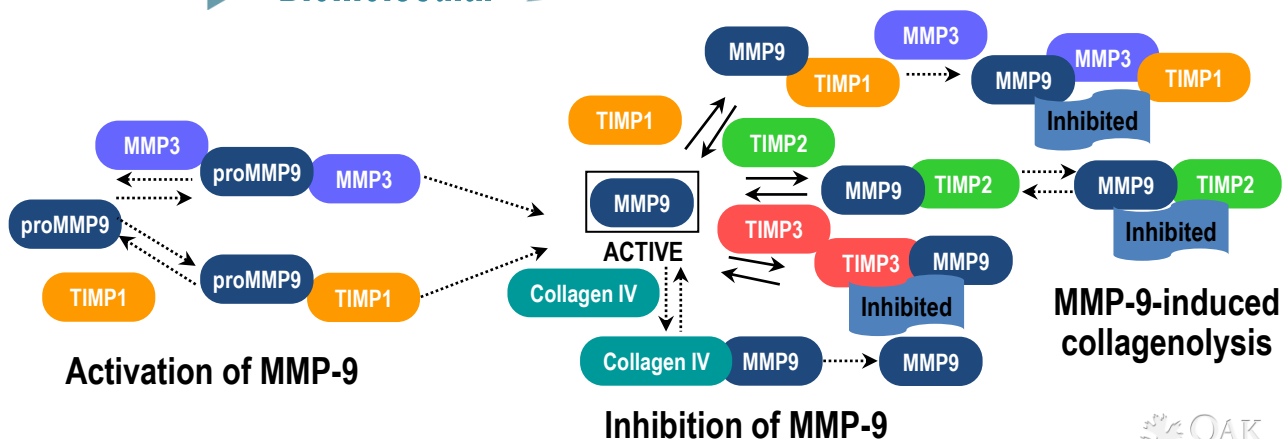
- Spatial modeling of cell migration
- Diffusive and kinetic modeling of biochemicals
- Result: A multiscale hybrid continuous-discrete predictive model for tissue pathology



Matrix metalloproteinases (MMPs)



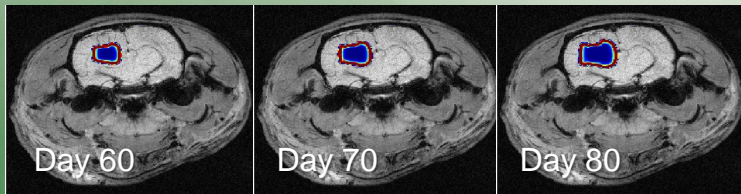
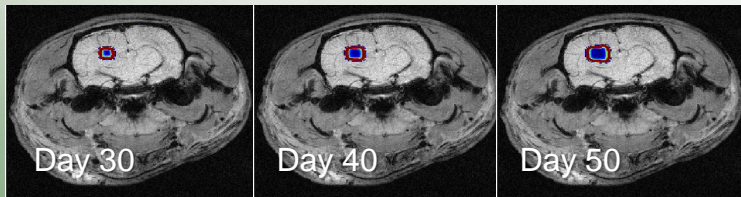
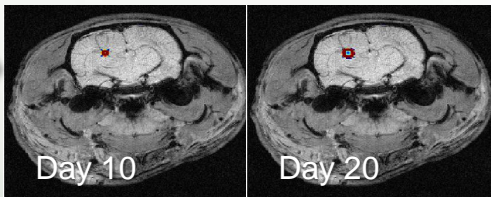
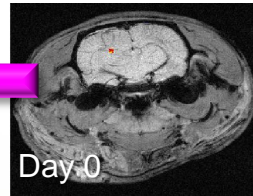
R. Conway 2005, zoologynews.blogfa.com/post-23.aspx



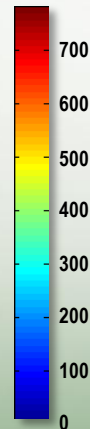
Qualitative system identification for tumor modeling

Modeling and Simulation Group

Computational Sciences & Engineering Division



Cells



Problem Statement:

ORNL is working with collaborators at the Vanderbilt University Institute of Imaging Science (VUIIS) to build and demonstrate a high performance, inductive reasoning engine for discovering models of tumor growth from features in time series images of mouse models of human breast cancer

Technical Approach:

Our approach integrates fuzzy inductive reasoning, genetic algorithms, and high performance computing to enable the construction of dynamic models *directly* from imaging characteristics that correlate with disease outcome

Benefit:

New methods and technology for the identification of clinically relevant, prognostic features that separate responding and nonresponding tumors early in the course of therapy, and a general and practical method for modeling biological processes directly from features in imaging data

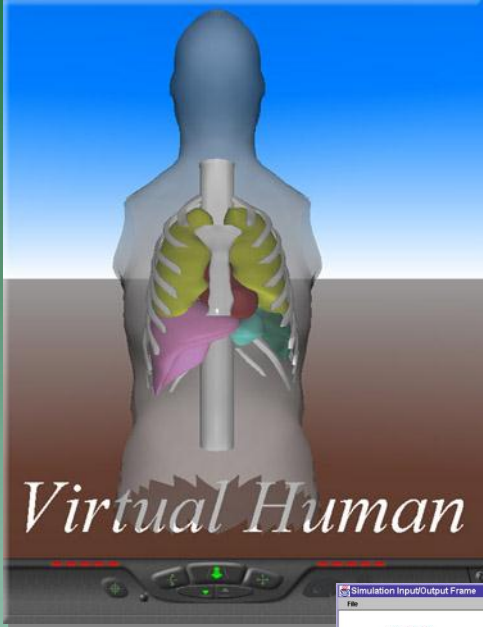
Point of Contact:

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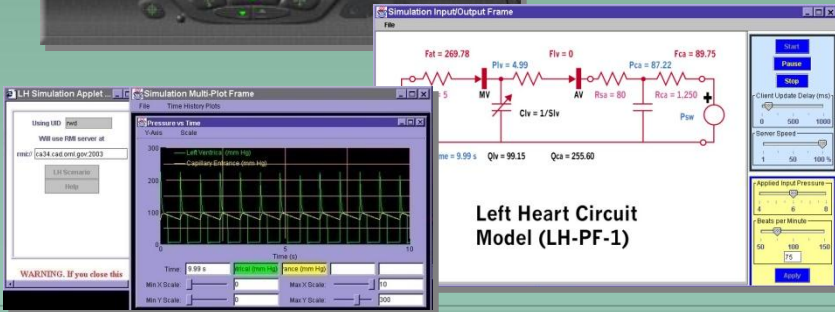


Cardiovascular modeling environments

Integrate

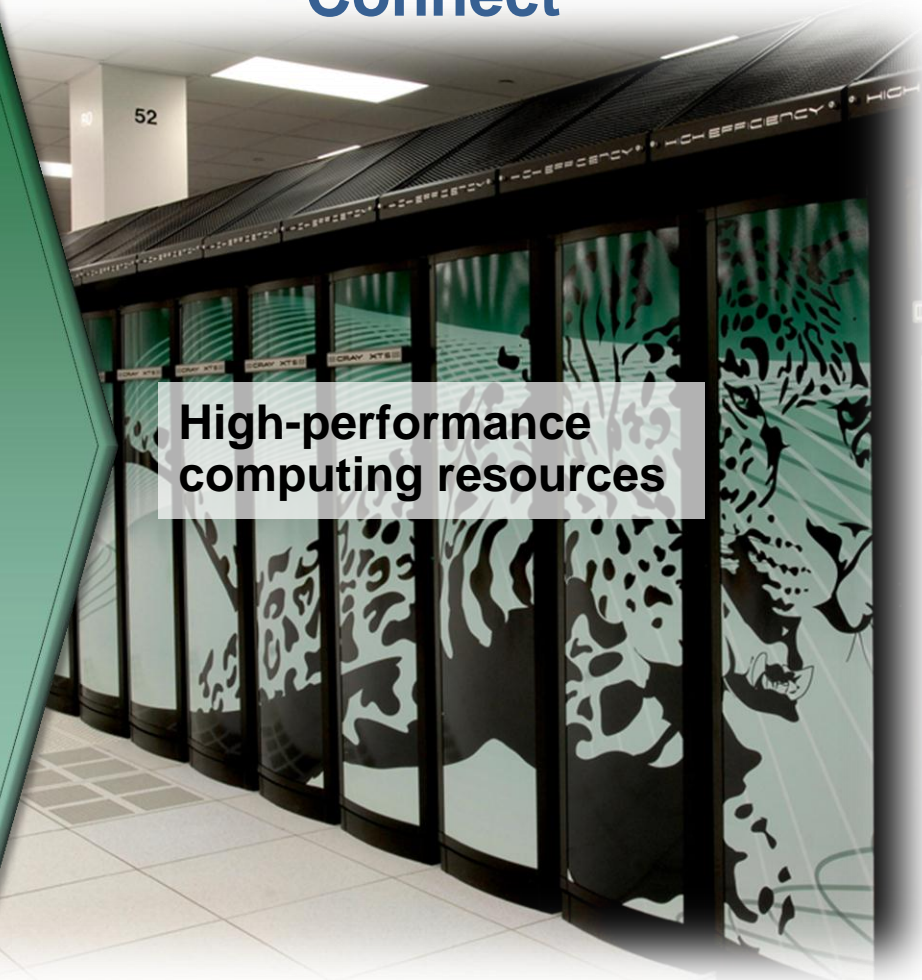


- Models
- Computations
- Visualization
- Predictions



Connect

High-performance computing resources



Neuroscience research areas



TBI

<http://www.brain-club.com/result.php?Keywords=acquired+brain+injury&host=www.brain-club.com&cat=1>

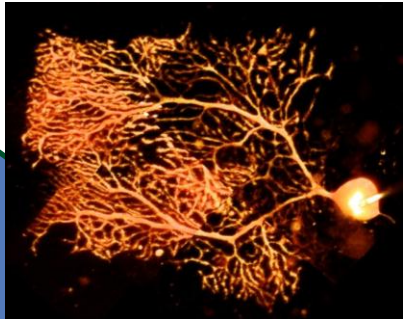


Image provided by Ryan Kerekes, ORNL

Neuronal Development/Migration



Nanostructured Materials

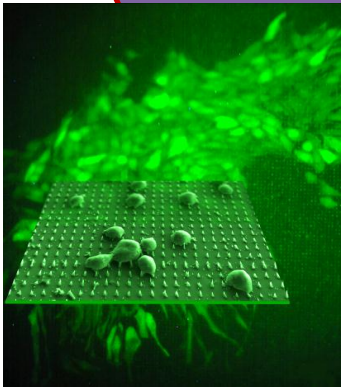
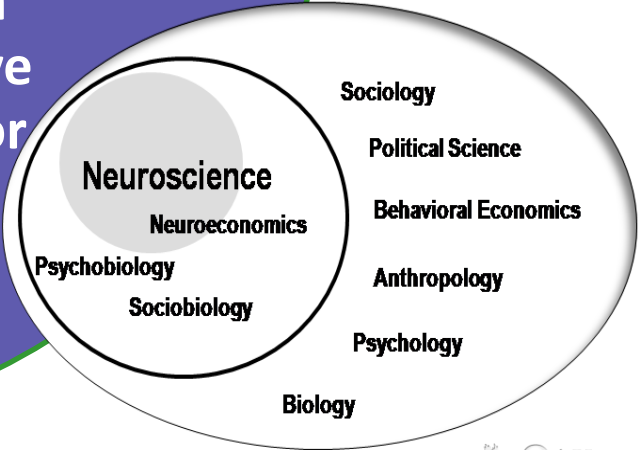


Image provided by Tim McKnight, ORNL

Human Cognitive Behavior



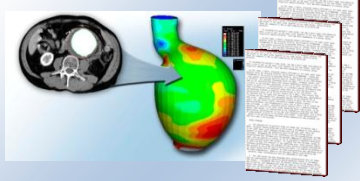
Multimodal data integration

MULTIMODAL DATA INTEGRATION WITH CROSS-DOMAIN APPLICABILITY

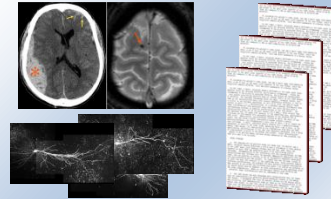
mammography data



aortic aneurysm studies

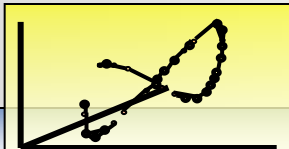


other domains

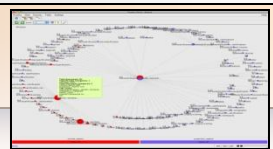


FRAMEWORK DEVELOPMENT

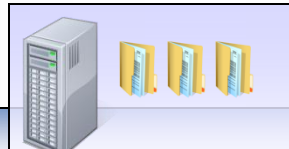
FUTURE WORK



multimodal manifold learning technique



clustering and data analysis



intelligent data retrieval system



large-scale data mining capabilities



USER-QUERYABLE DATA MANAGEMENT TOOL

Validation by Researchers and Clinicians

DAMSEL: Data Analytics for Medicine Using SEmi-supervised Learning*

Objective:

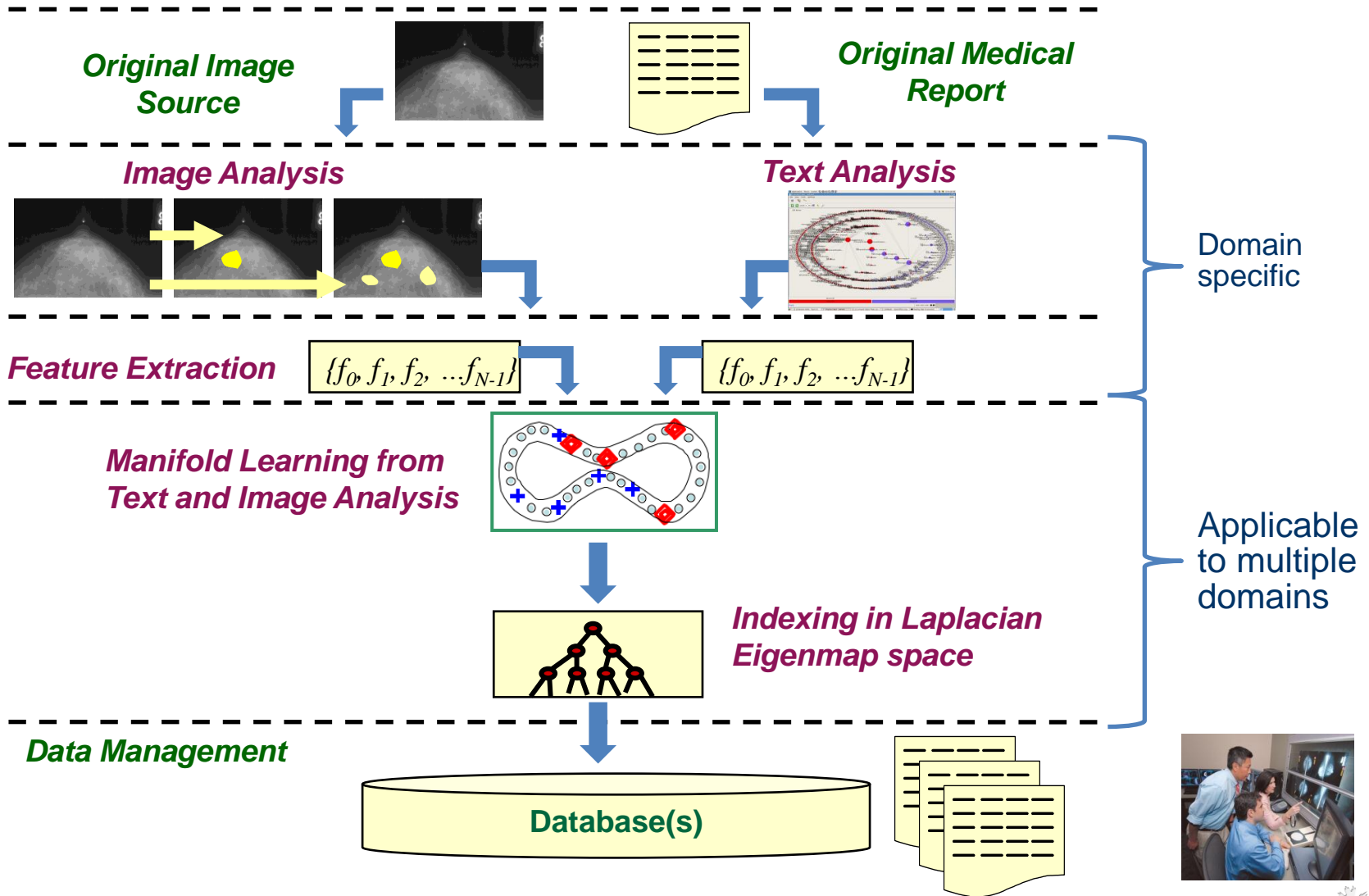
Develop a comprehensive, flexible, and responsive computing framework that incorporates ability to analyze multimodal data using semi-automated learning environments

Benefit:

Organized, integrated, feature-specific focus on relevant multimodal data for more effective, targeted analysis and decision making

* Research sponsored by the Laboratory Directed Research and Development Program of Oak Ridge National Laboratory, managed by UT-Battelle, LLC, for the U.S. Department of Energy.

DAMSEL concept: Text and image analysis in machine learning environment



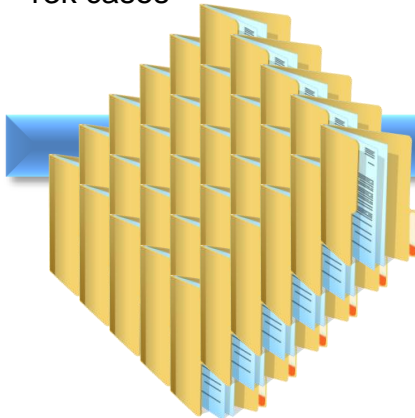
Text processing and features extraction

Objectives

- Leverage *human expertise* to characterize and classify the data
- Enhance statistical analysis with evolutionary algorithms to learn domain-specific, significant textual features
- Develop new classification learning algorithm based on evolutionary algorithms
- Provide basis for longitudinal analysis in order to find trends or precursors

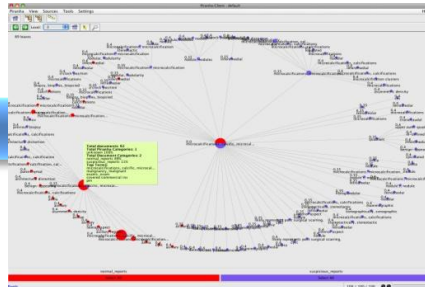
Database

- Text reports provided by the University of Chicago
- Total: 61k reports for 13k cases



Initial processing

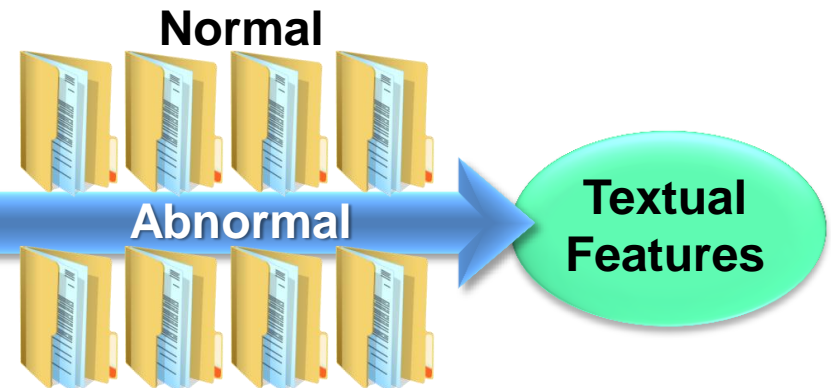
- Stopword removal
- Phrase extraction



Features extracted

- “No mass lesions evident”
- “Appearance of soft tissue densities”
- “Atypically located intramammary lymph node”

Statistical Analysis and Classification Learning Using Evolutionary Algorithms



Features extracted

- **Normal Reports:** “no & malignancy,” “no & masses”
- **Abnormal Reports:** “core & biopsy,” “spot & magnification”

Image processing and features extraction

Objectives

- Select suitable image processing algorithms to perform feature extraction
- Follow the Breast Imaging-Reporting and Data System (BI-RADS) specifications, and develop one scalable algorithm for each feature it describes
- Develop new techniques around the Fractal Segmentation concept
- Implement the algorithms on GPU platform toward large database processing

Database

- Images provided by the University of Chicago
- Total: 54k images for 13k cases

Initial processing

- Tissue segmentation
- Morphological assessment

Calcification, Mass, and Architectural Distortion Detection

Features extracted

- **Calcification:** number / distribution / contrast / cluster size / etc.
- **Mass:** shape / margin / density / texture / etc.
- **Architectural Distortion:** texture distortion

Features extracted

- Number of images (CC & MLO)
- Breast tissue and pectoral muscle segmentation
- Dimensions, asymmetry, texture, etc.

Images
Features

DAMSEL biomedical framework—multimodal manifold discovery

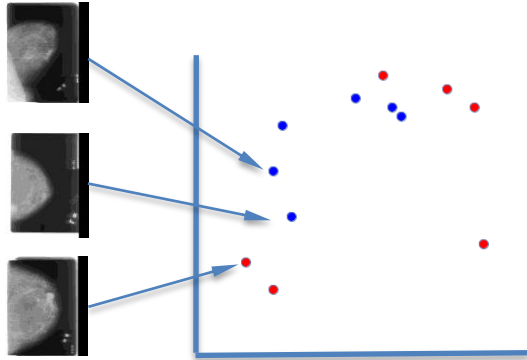
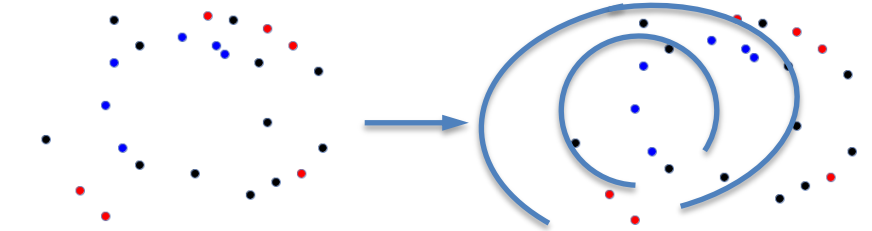
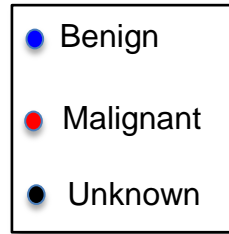


Image Space



Unclassified Points Help Reveal Manifold

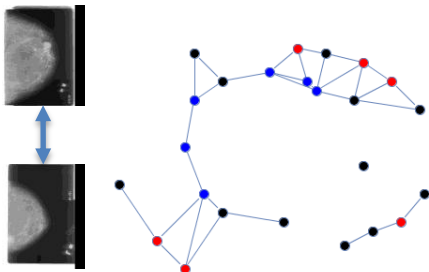
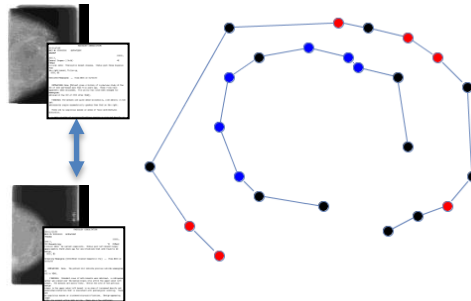
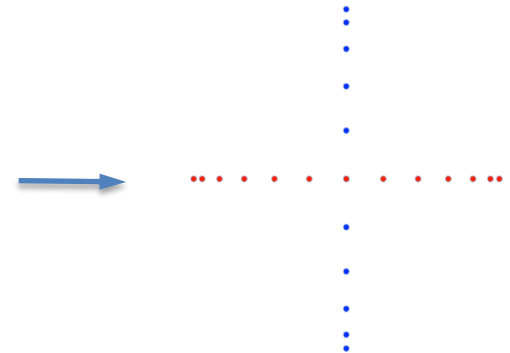


Image Similarity Graph



Similarity Augmented By
Cancer-Relevant Text

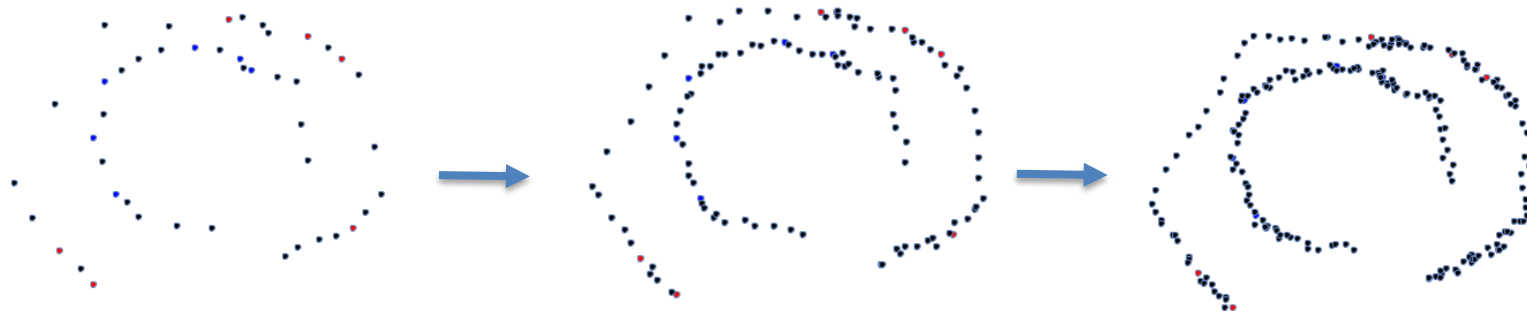


Improved Graph Laplacian-
Based Decision Space

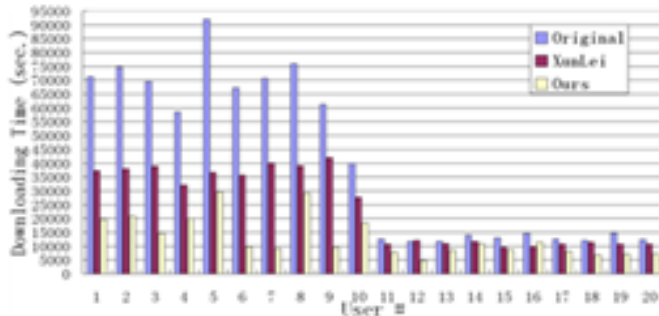
Increasing manifold fidelity

Attempts to significantly increase fidelity can impose great computational demands

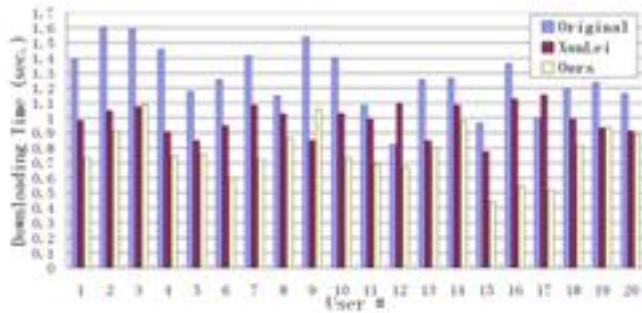
| Increasing: | Increases Complexity: | | |
|---|-----------------------|-------|--------|
| | Time | Space | Sample |
| Image/text feature set size | ↑ | ↑ | ↑ |
| Image/text feature complexity | ↑ | — | — |
| Patient sample set size | ↑ | ↑ | N/A |
| Thresholded, unweighted sparse graph → weighted dense graph | ↑ | ↑ | N/A |



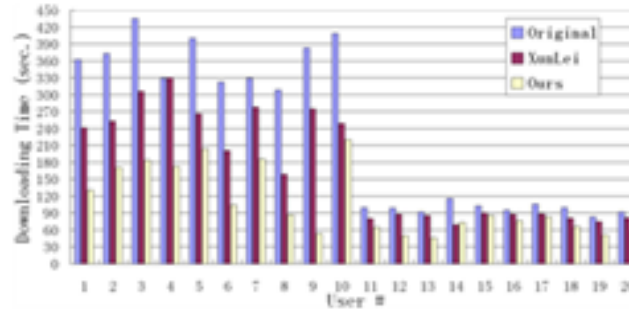
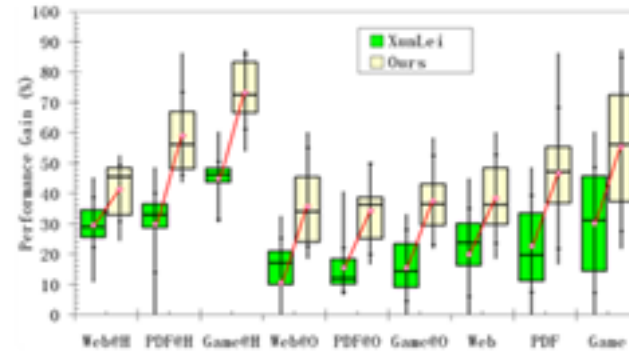
Multimodality data fusion



Game (784M)



Web (10.6K)



PDF (3.49M)

| Keyword Extraction Method | Precision | Recall | F-rate |
|---------------------------|--------------|--------------|--------------|
| TF × IDF | 0.210 | 0.312 | 0.251 |
| Yahoo! Term Extraction | 0.231 | 0.362 | 0.282 |
| Wikify! | 0.285 | 0.421 | 0.340 |
| Community detection | 0.312 | 0.435 | 0.373 |
| Our method | 0.456 | 0.513 | 0.483 |

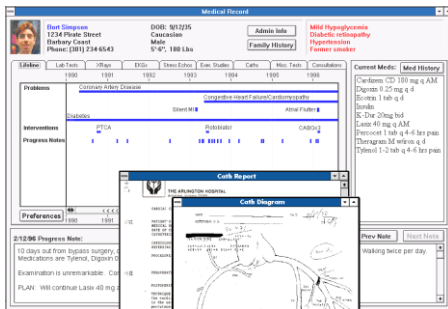
Slide provided by Songhua Xu

Computer tools for medical analytics and visualization (combine M&S, KD, and Viz)

Patient-centered care

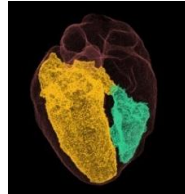


Patient medical record



Patient record

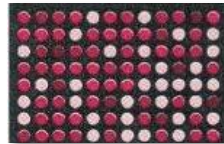
Data sources



Imagery

Tissue biopsy

Microarray data



Databases/system architecture

Advanced analysis and visualization

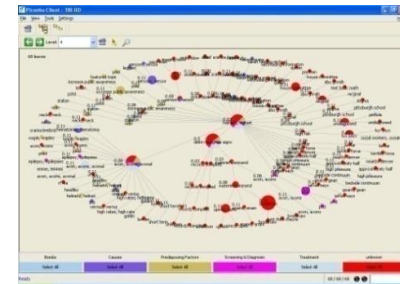
Image retrieval, processing and integration

Visualization

Knowledge discovery engine

Data analytics

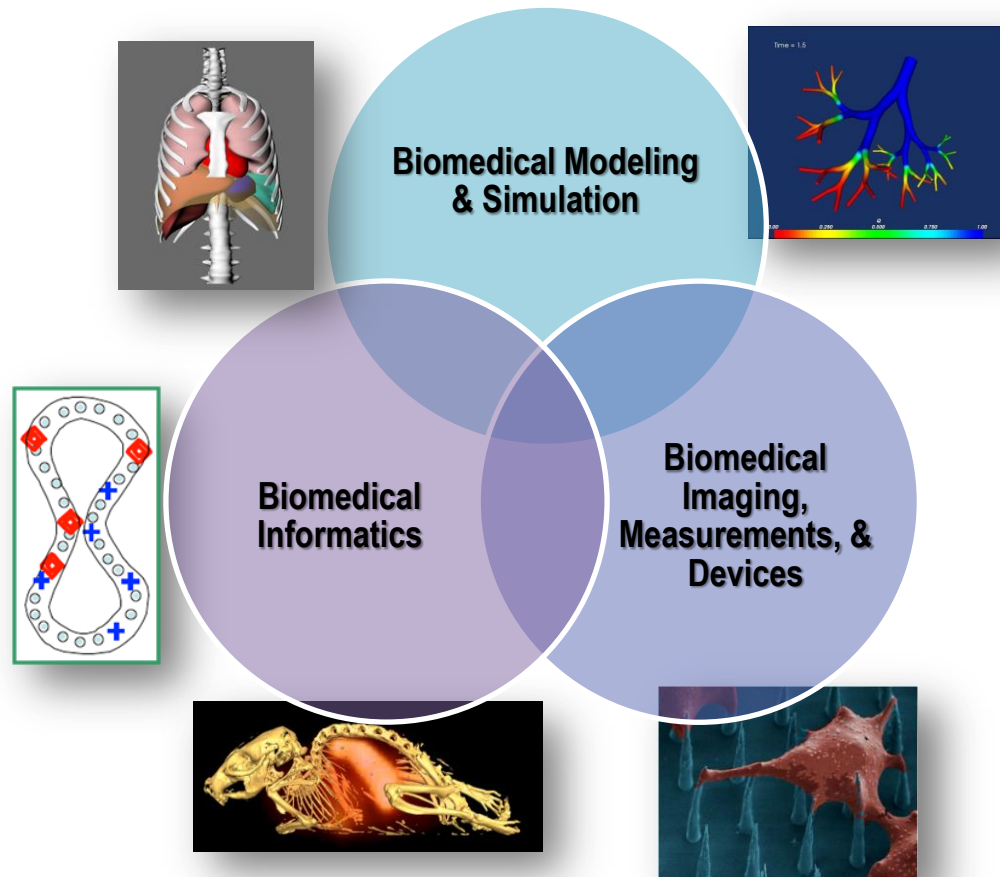
System architecture integrates databases



KD output

Biomedical Science and Engineering Center

Mission: To act as a catalyst and national resource for interdisciplinary biomedical research addressing challenges in key areas:



OBJECTIVES

- Establish a framework to identify critical research directions and facilitate interdisciplinary collaborations between academia, government agencies, the medical community, industry, and the ORNL research community
- Foster innovation and vital technological breakthroughs in key areas of the biomedical sciences
- Facilitate and organize an annual conference, providing a platform for interactions with nationally recognized biomedical science and engineering experts. Outputs from this conference will serve as inputs into roadmapping activities at NIH, DOD, and other medically related entities/efforts
- Establish and provide mentoring and educational opportunities for minority and underrepresented communities

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BSEC key activities

Conferences

1st Annual BSEC
Conference, March
18–19, 2009 at ORNL

http://computing.ornl.gov/cse_home/cms/conference/index.shtml

BSEC 2010,
May 25–26, 2010
Oak Ridge, TN

https://www.ornl.gov/bsec_conferences/2010/Index.shtml

BSEC 2011
March 16–17, 2011
Knoxville, TN

Seminars

Nationally and
internationally
known speakers

BSEC members

Education

Capstone Design
Program (Joint
BSEC/ORNL/UTK
program)

- Student internships
- Mentoring program
- Joint appointments (faculty)
- Core universities

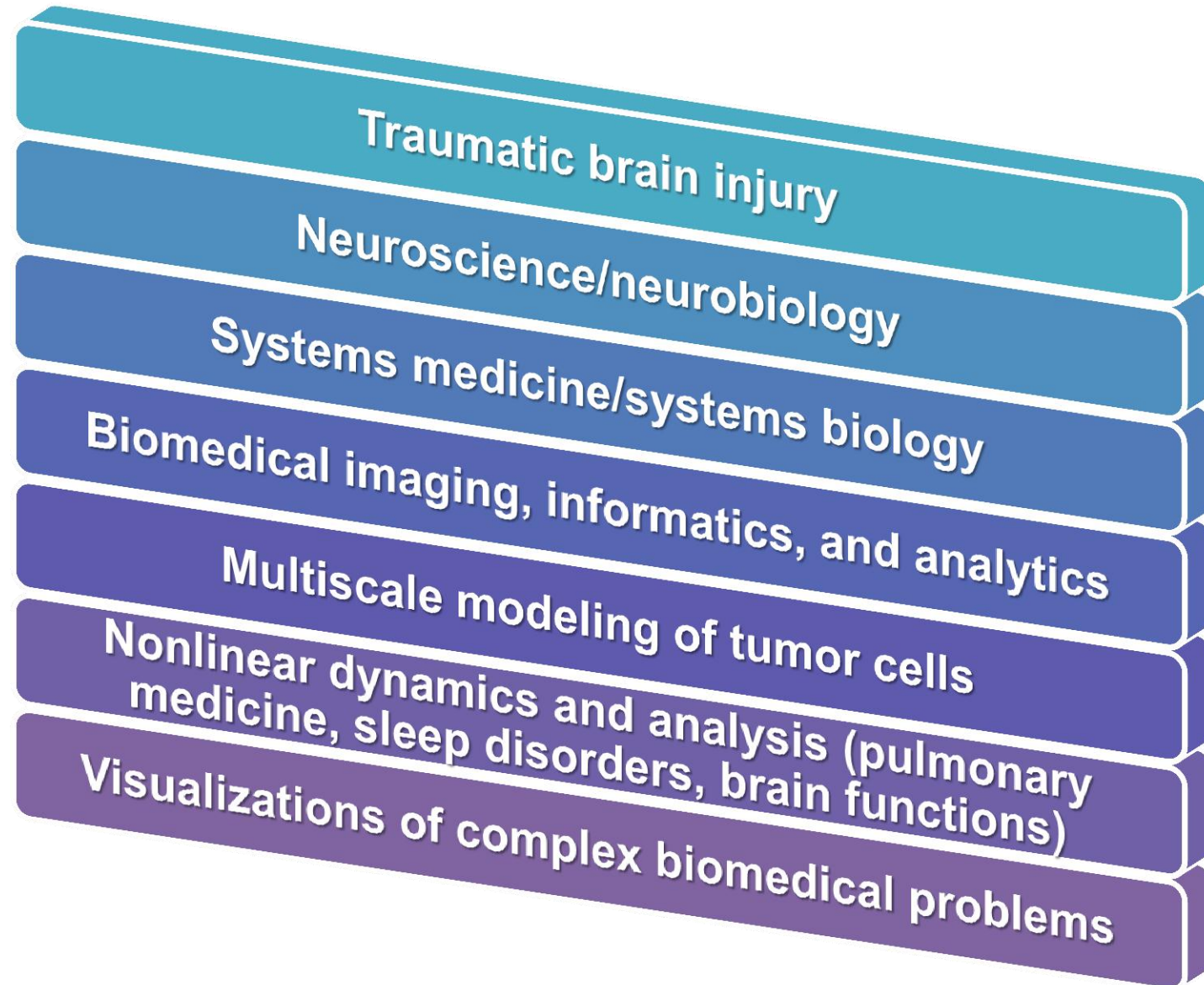
Collaborations and Partnerships

Universities

Federal agencies

Private sector

Growth opportunities and future directions



Contact

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