

# Johns Hopkins Pathology Informatics

- Joel Saltz, MD, PhD - Director
  - Support to Johns Hopkins Pathology
  - Research in Medical Informatics
  - Research in Computer Science
  - Development of External Software Products
  - Management Consulting
  - Partnership with Computer Science Department, University of Maryland College Park
    - substantial ongoing externally funded applications-driven computer science research program

# JHMI/UMCP

## Informatics/Computer Science

### Research Program

- Joel Saltz
- Merwyn Taylor
- Jerry Rottman
- Alan Sussman
- John Davis
- Kilian Stoffel
- Mike Beynon
- Renato Ferriera
- Asmara Afework
- Charlie Chang

# Pathology Informatics

- Pathology Data Systems
  - M based laboratory information system
- Data Repository
  - Relational, object-relational database and knowledge base systems
  - Supports ad-hoc queries, data mining, quality control, decision support
- Pathology image management
  - Telepathology, digital archiving of anatomic pathology imagery, electrophoresis data, flow cytometric, DNA sequencing and cytogenetic data

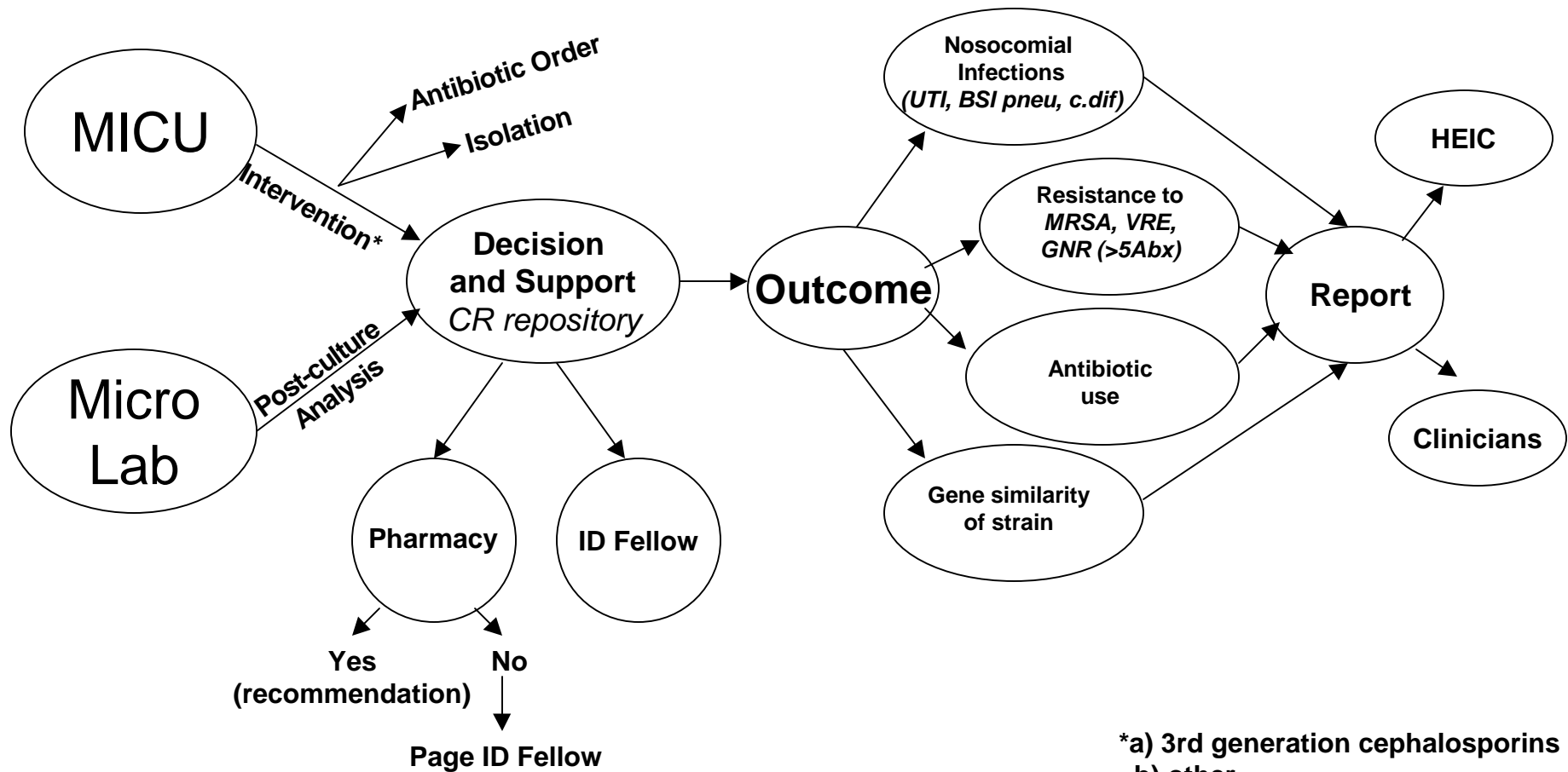
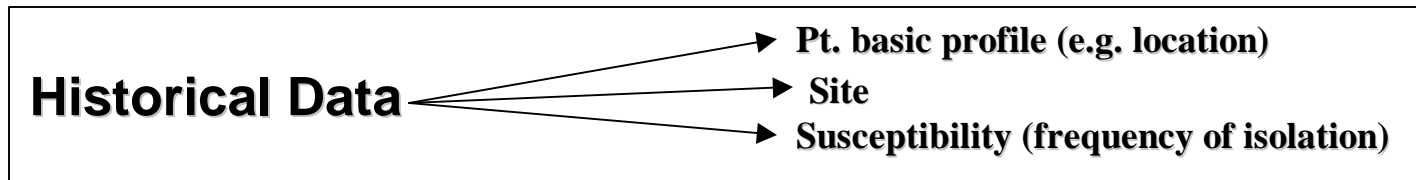
# Pathology Informatics

- Support for generation of interpretations
  - Support for interpretation of laboratory tests and for surgical pathology
  - Marshal data from LIS, electronic patient records, SPEP and UPEP results, surgical pathology imagery
  - Develop rules to generate a “first pass” interpretation
- Management Consulting
  - Combine data from multiple sources to analyze workflow, costs
- Research
  - Externally funded research that leads to new products and to new concepts in computer science

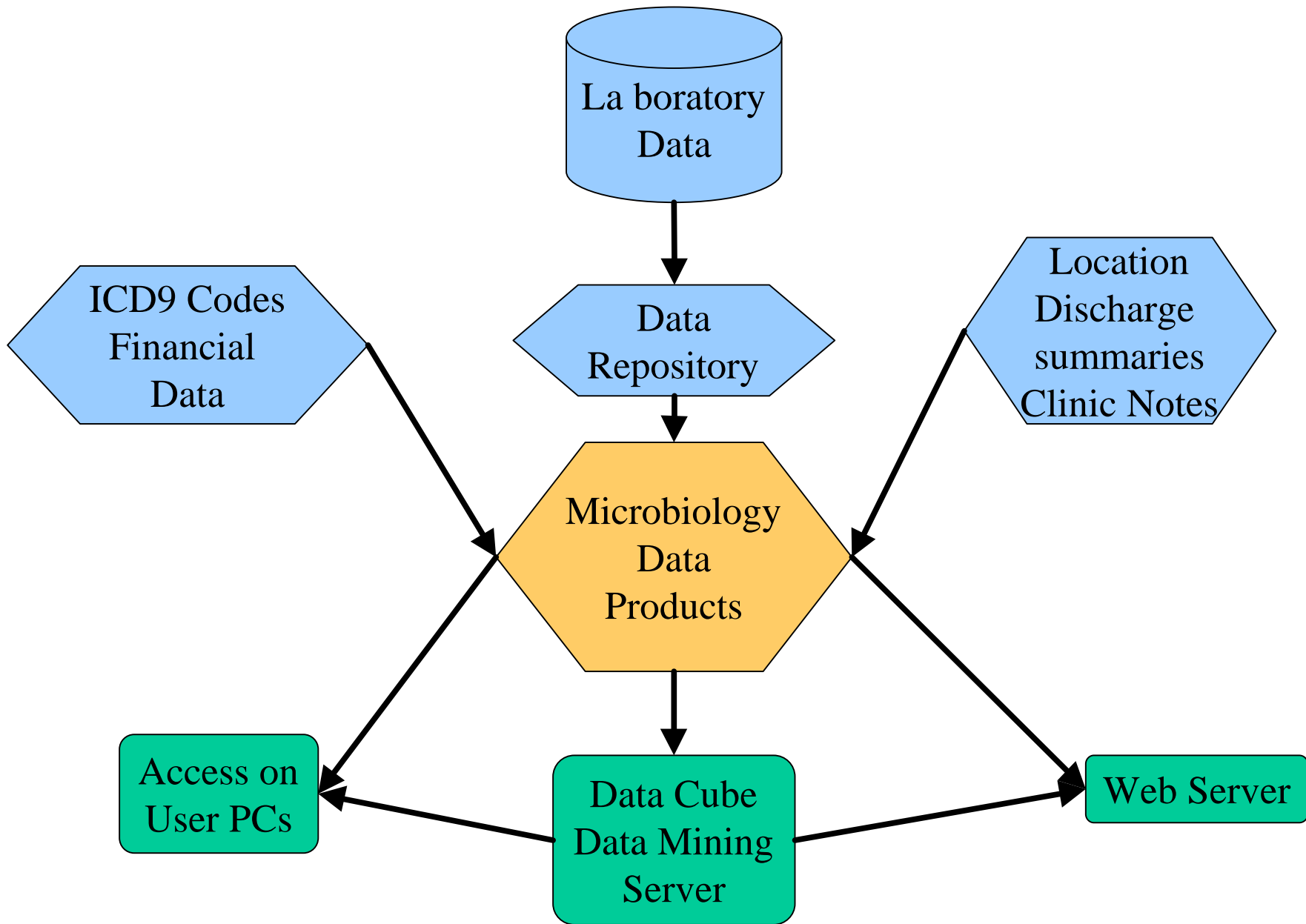
# Applications

- Management of Patient Care : *Gather and synthesize laboratory, clinical, cost and reimbursement data from many information systems to carry out long term management of large patient populations*
  - Antibiotic use and infection control (in development)
    - software to track and categorize infections
    - brings together laboratory, clinical and pharmacy data
    - flags unsuitable antibiotic use
    - flags antibiotic usage patterns that may lead to development of antibiotic resistance
    - screens for hospital acquired infections
      - potential for reducing antibiotic costs and for increasing accuracy of medical coding (with consequent increased hospital revenues)

# CDC Nosocomial Infection Pilot Plan



\*a) 3rd generation cephalosporins  
b) other



# Management of Patient Care

- Near Patient Testing (in development)
  - capture of patient laboratory data, combine with clinical data and intervene when clinically indicated
  - identify patients who would be appropriate candidates for new medical services
- Blood product use
  - evaluation of platelet use, response and reactions to transfusion therapy



# Applications

- Coordinated support for distributed laboratories: software to coordinate operation of laboratory and point of care laboratory testing devices (in development)
  - common management of data from point of care and laboratory instruments
  - ongoing assessment of quality of test results and of operator competence
  - use of all available data in quality assessments
  - real time feedback to prevent errors and to flag critical values
  - wedge for management consulting activities
    - assessment and reorganization of laboratory and near patient testing activities
  - interface with laboratory information systems

# Applications

- Dynamic digital telemicroscopy (in development)
  - software that allows local and remote users to capture, view, annotate and manipulate microscopy datasets
  - crucial addition to working medical record
  - facilitates comparison with other specimens from same and other patients
  - computerized medical education, conferences that involve multiple sites, certification, platform to run algorithms that grade tumors using morphometry, platform for pattern recognition algorithms
  - market would include both clients for viewing microscopy data and servers for storing and managing data

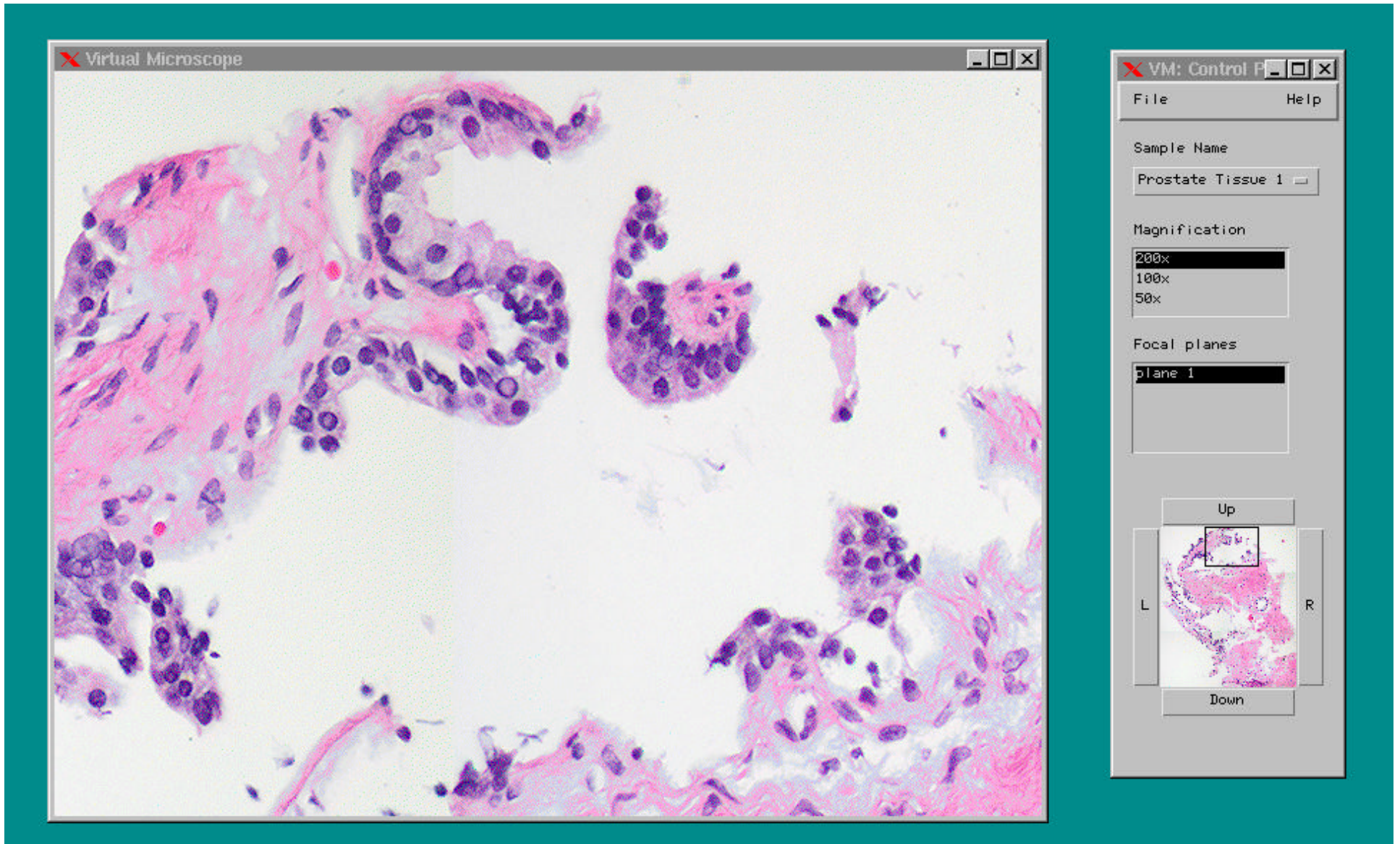
# Virtual Microscope

- World wide access to global collection of full digitized cases
  - no need to rely on slides available at local institution
  - compare specimens with earlier cases
- Research community can explore significance of morphology
  - compositing and 3-D reconstruction
  - quantitative immunohistochemistry analysis
  - run analysis programs on common data sets
- Links to relevant portions of digitized cases from medical information systems, electronic publications and textbooks.

# Training and Conference Environment

- Collection of institutions can carry out didactic conferences or collaborate on diagnosis
- Needle in haystack training
  - crucial skill often involves locating portions of case with interesting findings
- Multiple sites can
  - extends behavior of microscope - *independently* cruise through virtual pathology cases
    - users can change focus, magnification and move virtual stage
    - can track expert's examination of slide
  - capture whole case exploration process,
    - support combined medical record, demographic, pathology queries

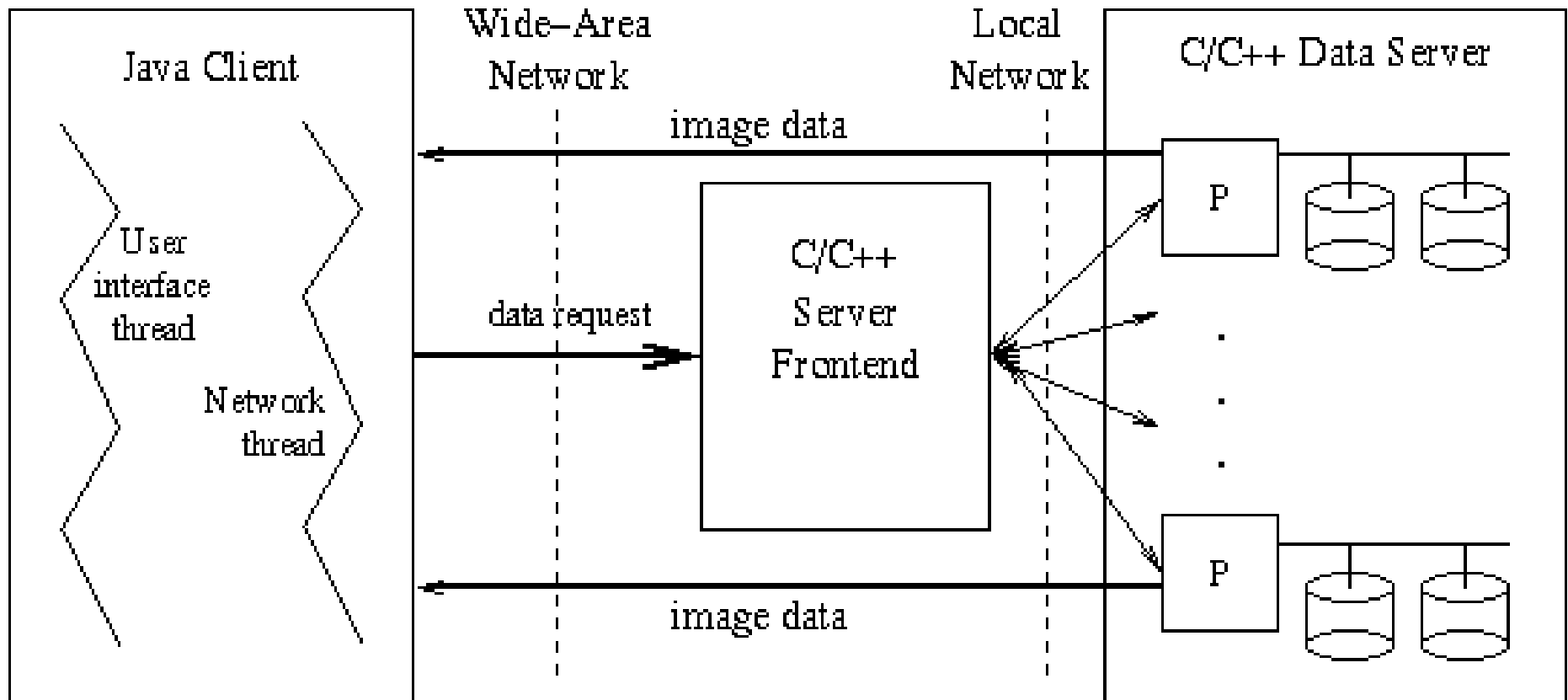
# Virtual Microscope Client



# Virtual Microscope Design

- Java based client
  - Client tested using Sun JDK and Microsoft J++
- Two part server (Currently runs on Windows, Solaris, AIX)
  - Front end -- accepts client queries, schedules queries and forwards to back end
  - Back end -- runs on multiple processors, retrieves and processes data from multiple disks

# Virtual Microscope Architecture



**Back end**

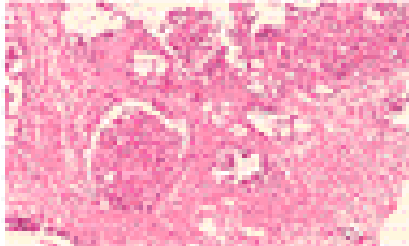


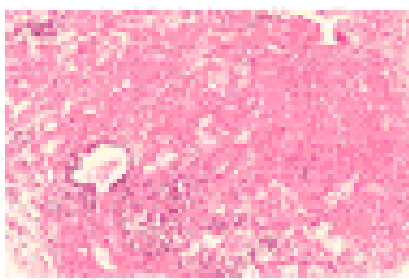
Virtual Microscope Test Slides - Microsoft Internet Explorer

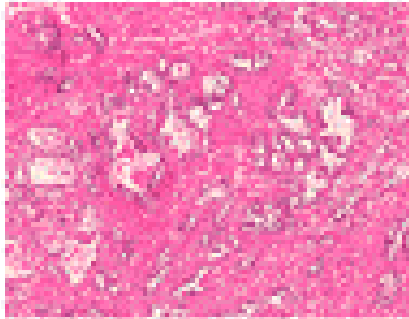
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Address <http://www.cs.umd.edu/users/rtfang/vmicrose/>

## Virtual Microscope Test Slides

 Breast tissue 1: (The diagnosis is atypical ductal epithelial hyperplasia). The clinical history is: The patient is a 63 year old woman with a prior history of total abdominal hysterectomy and bilateral salpingo-oophorectomy for well-differentiated endometrial carcinoma and benign breast cysts who is found to have focally clustered macrocalcifications on routine mammography. An incisional biopsy is performed. The attached slide is a representative section of this patient's histology.

 Breast tissue 2: (The diagnosis is non-atypical ductal epithelial hyperplasia (papillomatous) arising in the setting of fibrocystic disease of the breast). The clinical history is: The patient is a 35 year old with no significant past medical history but a family history of breast and colon cancer in first degree relatives who noticed a rubbery firm mass in the right breast on routine self examination. The patient underwent incisional biopsy of a 2.5 cm. mass, histology showed fibrocystic changes (cysts, apocrine metaplasia, stromal fibrosis) in most sections. One section is submitted for your review.

 Prostate tissue 2: (The diagnosis is adenocarcinoma of the prostate, Gleason grade 3+3=6, and background benign prostatic hyperplasia). The clinical history is: The patient is a 68 year old man with a prior history of colorectal adenomas, diverticulitis for which he underwent a left hemicolectomy, and several basal cell carcinomas of the skin and a family history of prostate carcinoma who presented with complaint of urge incontinence, nocturia, frequency, diminished force of his urinary stream and occasional dysuria. A screening prostatic specific antigen assay performed three months prior to the current evaluation was 5.5 (ng/ml); subsequent repeat biopsies showed no tumor.

Internet zone



# Data Mining and On line Analytical Processing

- On-line analytical processing tools to support queries that involve laboratory, pharmacy, clinical and financial data
  - complex hierarchies of categories
    - antibiotics, organisms, laboratory tests by particular test, clinical category, panel, test instrument
  - use hierarchies to formulate queries
    - screening for hospital acquired infections
    - targeting antibiotic use to optimize efficacy, reduce development of resistant organism strains, reduce costs
  - data cube and data mining software

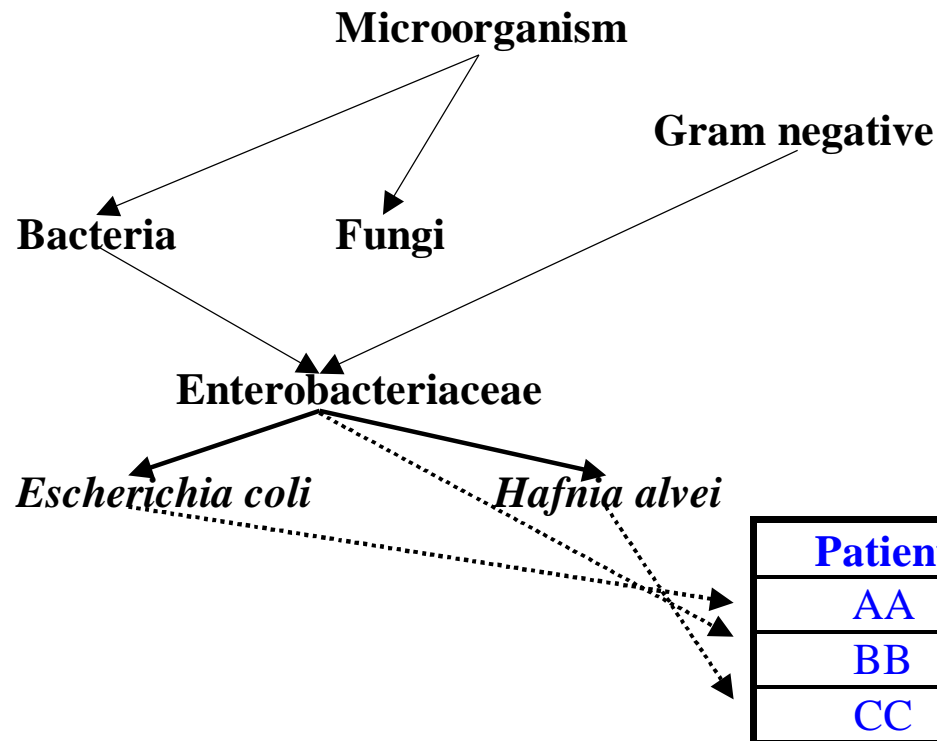
# Microbiology Queries

- For all classes of aerobic bacteria (ranging from general categories like gram negative rods to specific organisms like *Yersinia enterocolitica*) characterize resistance to all beta lactam antibiotics
- Compare effectiveness of different possible antibiotic protocols for neutropenic oncology patients
  - e.g. Trimethoprim/Sulfa + piperacillin v.s. Trimethoprim/Sulfa + piperacillin/tazobactam
- Screen for changes over time in antibiotic susceptibility

# Hierarchies and Databases

- Domain specific knowledge represented by hierarchies
- Graphical user interface or programmer API (currently two different versions) make it possible for the user to select portions of a hierarchy
- User can use a hierarchy to select a patient subset
  - user can then use other hierarchies to carry out data cube operation
  - e.g. for oncology patients, tabulate third generation cephalosporin antibiotic resistance in all classes of non-fermenter microorganisms

# Relationship between an Ontology and a Database



<b>Patient</b>	<b>Microorganism</b>	<b>Age</b>
AA	<i>Escherichia coli</i>	34
BB	<i>Enterobacteriaceae</i>	52
CC	<i>Hafnia alvei</i>	19

**DataBrowser**

Add to Query  
Clear Query  
Send Query

AND  
 OR

Start Date   
End Date

DATE

- ▶ Locations
- ▶ MICs
- ▼ Organisms
  - ▼ bacteria
    - ▼ bacterial
      - ▼ aerobe
        - ▶ gp aerobe
        - ▼ gn aerobe
          - ▼ nr aerobe
            - ▶ ebact
            - ▼ vibriona
              - ▶ AEROMONAS SPECIES
              - ▶ VIBRIO SPECIES
              - PLESEOMONAS SHIGELLOIDES

```
[[  
Location IN ('373CL','379CL','393CL')  
OR (  
Location IN ('ONC','ONC-2','OC-N3','GSONC','OC-S  
AND
```

**DataBrowser**

AND  
 OR

Start Date:   DATE  
 End Date:

- ▶ tetracyclines
- ▶ macrolides
- ▼ beta lactam
  - ▼ penicillins
    - ▶ penicillinase resistant pen
    - ▶ extended spectrum pen
    - ▼ penicillin inhibitor combination
      - ▼ pip/tazobactam
        - ▼ A\_P/T
          - ▶ P/T susceptible
          - ▶ P/T intermediat
          - ▶ P/T resisant
    - ▶ cephalosporins
    - carbepenems
    - monobactams

```

AND((((
  Field4 IN ('ebact','ESCH','ESCO','E0157'
  Field4 IN ('nonfermenter','PSEU','PSAE','
Field17 IN ('T_S_ENF_R','2+/38+'))))
OR ((
  Field4 IN ('PSEU','PSAE','PSAL','PSFL','F
Field17 IN ('T_S_PS_R','2+/38+'))))
AND((((
  Field4 IN ('ebact','ESCH','ESCO','E0157'
  Field4 IN ('nonfermenter','PSEU','PSAE','
Field21 IN ('P_T_ENF_R','64+/4+'))))
OR ((
  Field4 IN ('PSEU','PSAE','PSAL','PSFL','F
Field21 IN ('P_T_PS_R','64+/4+'))))
  
```

Cube Interface

nr\_aerobe [S | R (total)]

ms

- acteria
  - [-] bacteria1
    - [-] aerobe
      - [+] gp\_aerobe
      - [-] gn\_aerobe
        - [-] nr\_aerobe
          - [+] ebact
          - [+] vibriona
          - [+] nonfermenter
          - [+] nr\_fast
        - [+] nc\_aerobe
        - [+] spiro
      - [+] mycoplasma
      - [+] afb
    - [+] anaerobe

antibiotics

- [+] aminoglycosides [81.01 6.92 12.06 (2412)]
- [+] quinolones [79.18 8.43 12.39 (759)]
- [+] tetracyclines [14.23 37.38 48.39 (808)]
- [+] macrolides [32.0 0.0 68.0 (25)]
- [+] glycopeptides [82.61 0.0 17.39 (23)]
- [-] beta lactam [42.1 14.15 43.75 (5772)]
  - [-] penicillins [32.61 15.88 51.51 (2551)]
    - [+] penicillinase resistant pen [52.0 0.0 48.0 (25)]
    - [+] extended spectrum pen [32.7 15.94 51.37 (2416)]
    - [+] penicillin inhibitor combination [26.36 18.18 55.46 (11)]
  - [+] cephalosporins [49.61 12.79 37.6 (3221)]
- [+] Trim/Sulfa [62.21 8.67 29.12 (807)]

Calculate Distribution

Calculate Distribution

Evaluate Cube

EXIT