

# Motivation

- Need to distribute results for review
- Create with multiple tools in different formats
- Reviewers without access to original tool
- Measurement technique comparisons
- Imaging bio-marker development/testing
- Image-based clinical trials, esp. oncology
- Change in measurements over time
- Clinical practice for individual patients

# Results Characteristics

- What is recorded?
  - measurements – distance, volume, density, etc.
  - coordinates – what region on image measured
- Context?
  - identification of subject (patient), lesion, etc.
  - identification of reader
  - identification of technique
  - position in time (change over time, no change)

# Results Organization

- Single object per measurement
- Single object for all measurements at one time
- Single object per reader per time point
- Single object per reader all time points
- Single object per image with all measurements for all readers
- ...

# Use-Cases

- QIBA CT Volumetry 1B Round 2
  - lung cancer volume measurement
  - multiple independent readers
  - two time points
  - some cases no change, some with change
  - volume & automatically derived distance
    - standard DICOM SR and DICOM Segmentation format
    - clinical trials results information model
    - organized as one object per lesion per reader
  - reading tool is not distributable for review

# Use-Cases

- QIBA CT Volumetry 1A
  - phantom lung nodules differing size and shape
  - multiple independent readers
  - one time point
  - two different (incompatible) reading tools used
  - distance
    - proprietary format
    - one object per image containing multiple readers & lesions
  - volume
    - variant of DICOM RT Structure Set
    - one object per reader per lesion
  - reading tools are not distributable for review

# Hierarchical Model

- Subject
  - Reader
    - Time Point
      - Lesion
        - » Region (-> link to image coords)
          - Measurement (e.g., Volume)

# Hierarchical Model

- Subject = 0001
  - Reader = 1
    - Time Point = 2010/06/01
      - Lesion = 1
        - » Volume = 355 mm<sup>3</sup>
      - Lesion = 2
        - » Volume = 3896 mm<sup>3</sup>
    - Time Point = 2011/07/01
      - Lesion = 1
        - » Volume = 471 mm<sup>3</sup>
      - Lesion = 2
        - » Volume = 3801 mm<sup>3</sup>
  - Reader = 2
    - ...

# One Annotation Per File

- Subject = 0001
  - Reader = 1
    - Time Point = 2010/06/01
      - Lesion = 1
        - » Volume = 355 mm<sup>3</sup>
      - Lesion = 2
        - » Volume = 3896 mm<sup>3</sup>
    - Time Point = 2011/07/01
      - Lesion = 1
        - » Volume = 471 mm<sup>3</sup>
      - Lesion = 2
        - » Volume = 3801 mm<sup>3</sup>
  - Reader = 2
    - ...



# One Time Point Per File

- Subject = 0001
  - Reader = 1
    - Time Point = 2010/06/01
      - Lesion = 1
        - » Volume = 355 mm<sup>3</sup>
      - Lesion = 2
        - » Volume = 3896 mm<sup>3</sup>
    - Time Point = 2011/07/01
      - Lesion = 1
        - » Volume = 471 mm<sup>3</sup>
      - Lesion = 2
        - » Volume = 3801 mm<sup>3</sup>
  - Reader = 2
    - ...

# One Reader per Subject Per File

- Subject = 0001
  - Reader = 1
    - Time Point = 2010/06/01
      - Lesion = 1
        - » Volume = 355 mm<sup>3</sup>
      - Lesion = 2
        - » Volume = 3896 mm<sup>3</sup>
    - Time Point = 2011/07/01
      - Lesion = 1
        - » Volume = 471 mm<sup>3</sup>
      - Lesion = 2
        - » Volume = 3801 mm<sup>3</sup>
  - Reader = 2
    - ...

# Tabular Presentation

Subject	Reader	Time Point	Lesion	Volume mm3
0001	Reader 1	2010/06/01	1	355
0001	Reader 2	2010/06/01	1	375
0001	Reader 1	2010/06/01	2	3896
0001	Reader 2	2010/06/01	2	4764
0001	Reader 1	2011/07/01	1	471
0001	Reader 2	2011/07/01	1	289
0001	Reader 1	2011/07/01	2	3801

# Tabular Presentation

- Can be sorted by different columns
- Easy to add derived computations
  - e.g., % difference from mean volume
- Exportable to spreadsheet tools (e.g., Excel)
- Exportable to statistical tools (e.g., R)
- Easy to feed selected columns to chart tools
- Cells and rows can be hyperlinked to images
  - e.g., hyperlink a volume to the ROI outline

# Structured vs. Unstructured

- Structured input defines individual elements
  - measurements are recorded
    - as opposed to recalculated from coordinates each time
  - measurements & units distinct & related to image
    - coordinates linked to measurement
    - different types of measurement coded (not free text)
  - individual context elements distinct
    - e.g., patient, lesion, reader encoded separately
- Unstructured annotations
  - user enters free text (e.g., “Reader 1 Lesion 1)
  - text and coordinates linked, or
  - text and coordinates co-located but not linked

# Results Formats

- DICOM Structured Reports
  - general purpose hierarchical data format
  - primitives for codes, measurements, coordinates
  - references to images, segmentations
  - needs a “template” to define information model
  - no widely adopted standard templates for oncology quantitative measurements over time
  - author has defined template for internal use in contract research, and has reused it for QIBA

# Results Formats

- DICOM Segmentations
  - rasterized (pixel array) of values matching image
  - encoding of lesions
    - Binary - which voxels are included in lesion
    - Probability maps – probability voxel is included
  - encoding of “label maps”
    - E.g., atlas of tissue types corresponding to voxels
  - does not contain measurements
    - intended to be referenced from a DICOM SR

# DICOM RT Structure Sets

- Developed for and long history of use with Radio-Therapy Planning applications
- Generic mechanism for 3D contours
  - set of coplanar iso-contours
  - 3D patient-relative coordinates
- Image co-ordinates
  - often 1:1 correspondence of 3 coordinate with original image slices and voxels (though not required)
- Measurements
  - very limited, if any, measurements in file
  - recalculated on re-loading
  - could be referenced from a DICOM SR (though unusual)



# DICOM Presentation States

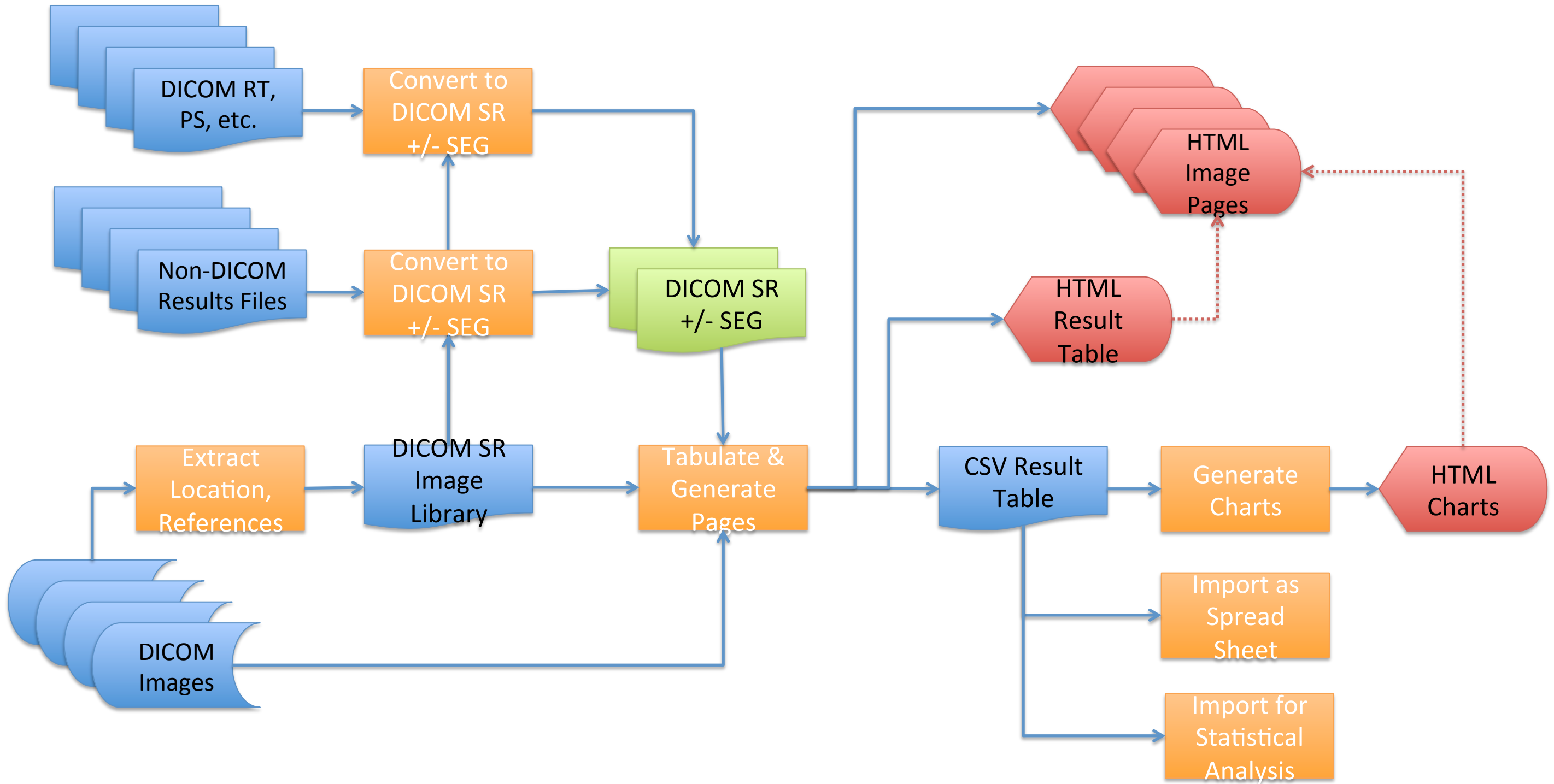
- Intended for rendering, not interpretation
- Commonly implemented in PACS for simple annotation capture
- Unstructured
  - text and graphics are not semantically linked
  - text is free text, not coded, and no structured measurements
  - with discipline entering text, structured content can be parsed from free text retrospectively

# Process Flow

- Create DICOM SR
  - that conform to a basic template
  - subject/reader/time point lesion/measurement
- Tabulate results
  - include hyperlinks to rendered images with ROIs
  - compute any derived statistics
- Generate charts from tables
  - e.g., scatter plots, waterfall plots

# Process Flow

- ROIs
  - if SR references segmentation, use it
  - if SR contains coordinates (2D or 3D), use them
  - if not, convert coordinates to SR coordinates
    - e.g., RT Structure Set 3D coordinates to SR 2D
  - if not, convert raster to segmentation
    - e.g., LIDC Max tool PMAP to DICOM SEG
- Image Library
  - extract image characteristics for re-use
  - e.g., position, orientation, spacing, UIDs
  - store in SR Image Library template
  - saves repeating this (reading image headers) many times

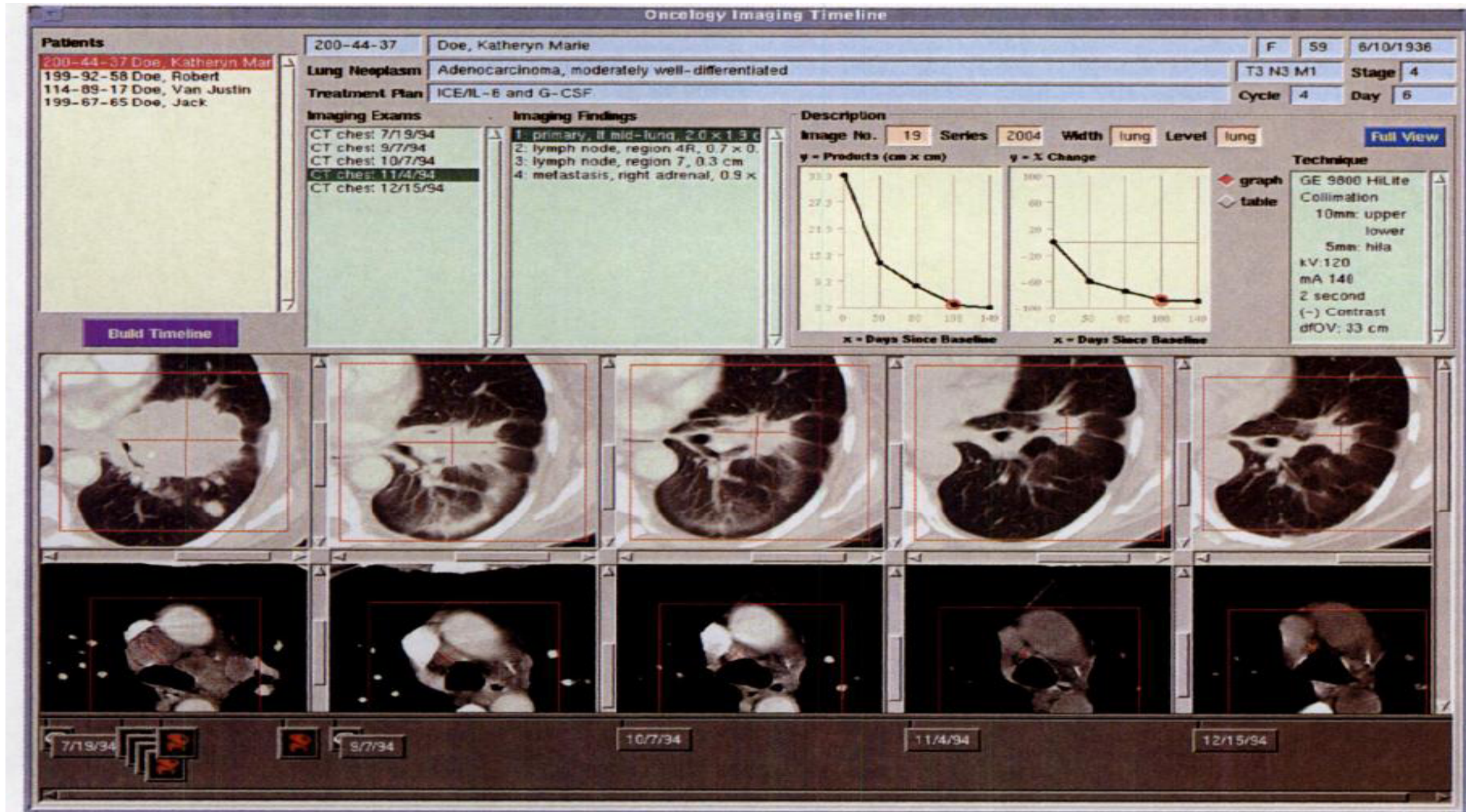


# Earlier Work by Others

- Aberle 1996 – Thoracic Oncology Imaging Timeline (OITL)
  - regions of interest defined during reporting
  - change in lesion size over time for single patient
  - table of lesion size
  - graphs of size change
  - visualization of size change

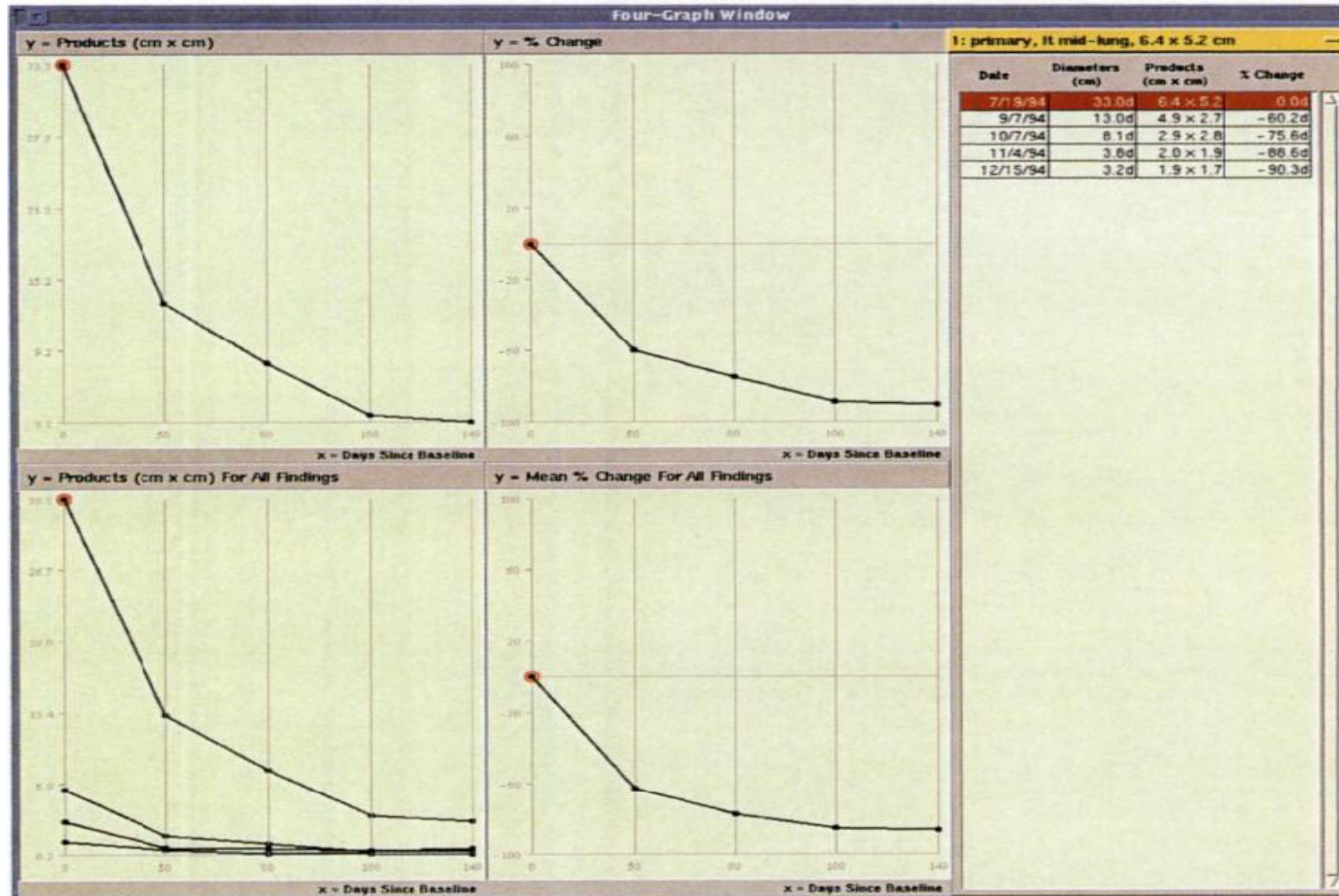
<http://radiographics.rsna.org/content/16/3/669>

# Aberle 1996 - OITL





# Aberle 1996 - OITL



# Earlier Work by Others

- Bui 2007 – TimeLine
  - more generalized, configurable approach
  - data access and integration
  - data mapping, reorganization and clustering
  - hierarchical problem-centric views
  - emphasis on temporal chronologies & clustering
  - adaptable format mapping methods

<http://dx.doi.org/10.1109/TITB.2006.884365>



# Bui 2007 - TimeLine

Patient Jane Doe  
Research ID 123-45-678

Integrated bone development report

May 16, 2003 (Baseline)      October 7, 2004 (Follow-up 1)      August 17, 2005 (Follow-up 2)

5/16/03 CT Scan L2

Children's Hospital of LA (CHLA\_000)

EMD

05/16/2003 10:26:42

slice thickness: 10.0 position: 110.58

pixel spacing: x = 0.674 y = 0.674

5/16/03 Integrated Data Panel

H&P		DEXA	Lumbar spine	Area	BMC	BMD	Reference curve
Age	19		L1	12.92	11.78	0.911	
Gender	F		L2	13.42	13.95	1.040	
Ethnicity	Lat		L3	15.37	16.20	1.054	
Tanner stage			L4	17.03	17.49	1.027	
Standing height	5'9"		Left hip				
Sitting height	3'2"		Right hip	Area	BMC	BMD	
Weight	132		Right hip neck	5.04	4.26	0.846	
BMI			Right hip troch	10.25	6.68	0.652	
Waist	29"		Right hip inter	24.28	26.77	1.103	
Hip	34"		Right hip Ward's	1.29	1.10	0.850	
Skeletal age	18		Whole body				

5/16/03 DE

XR DEXA BONE DUAL ENERGY

Baseline Exam. The average bone mineral density from L1-L4 is 1.011 g/cm<sup>2</sup>, 0.15 standard deviations below the mean for age and sex-matched controls. Compared to normal young adults, this density is 2.94 standard deviations below the mean or 69% of normal. The bone mineral density in the right total hip is 0.953. Compared to normal young adults, this density is 2.99 standard deviations below the mean or 61% of normal.

Normal bone mineral density of the lumbar spine. There is no age and sex-matched data for the hip.

Highlight NLP results for DEXA report

Show DEXA images in separate window

OK

Area     BMD     PR     T-score

BMC     T-score     Z-score   

Open DEXA report    Compare    OK

Search

Access BBN

Patient Database

# Earlier Work by Others

- Levy 2007 – LesionViewer
  - serial oncology studies
  - anatomical summary of lesion location
  - direct navigation to visualization of location
  - temporal abstraction of lesion behavior

<http://dx.doi.org/10.1109/TITB.2006.884365>

# Levy 2007 - LesionViewer

[View Scan Images](#)

[View Disease Summary](#)

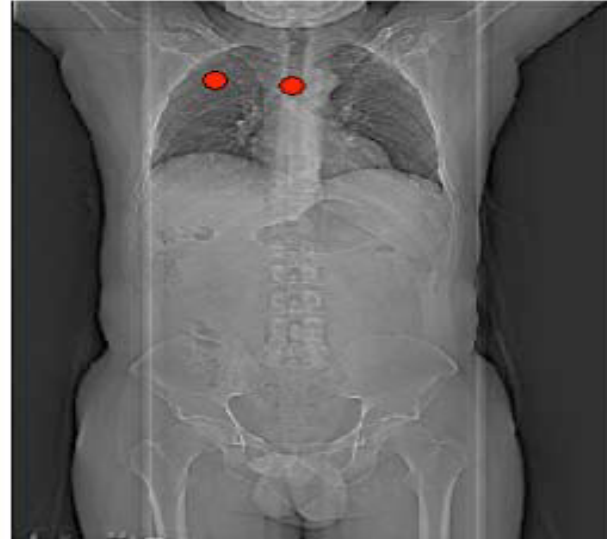
[View Raw Data](#)

[Select New Patient](#)

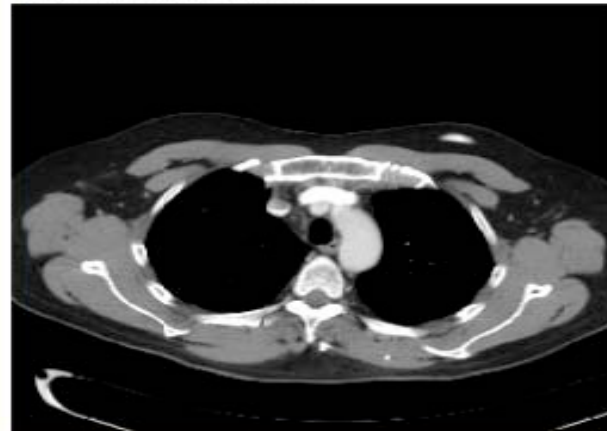
Name: Wilson, Mark  
MR Number: 55555555

Date: 03/28/06

CT Chest, Abdomen and Pelvis

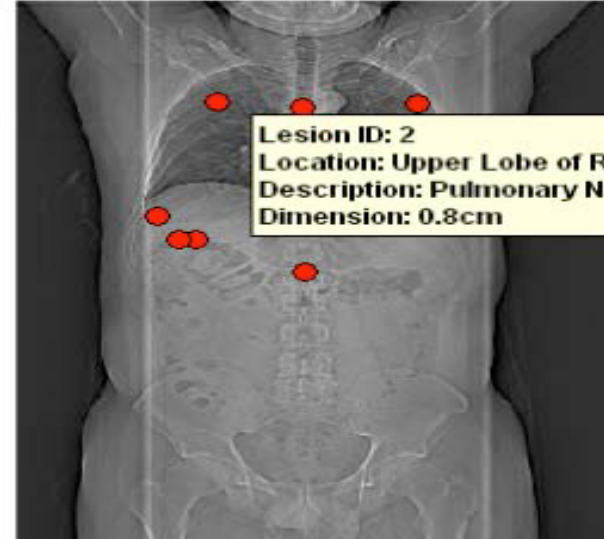


Pulmonary Nodule  
Dimension: 0.7cm

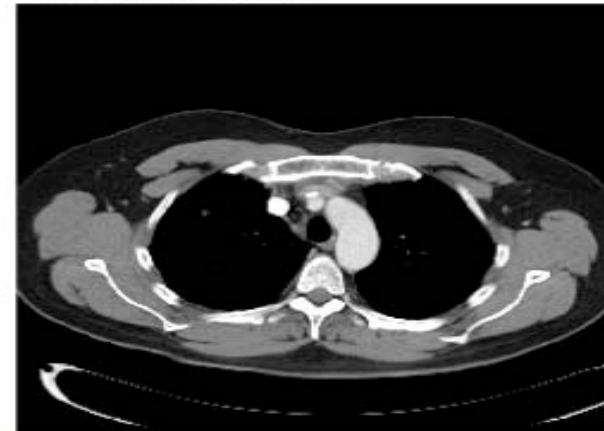


Date: 08/3/06

CT Chest, Abdomen and Pelvis

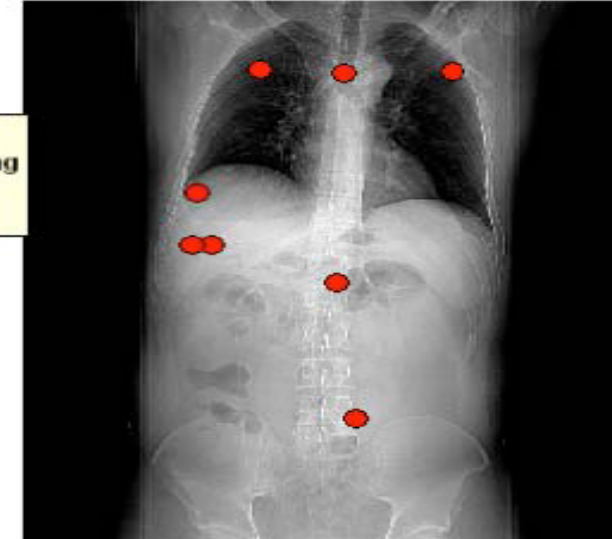


Pulmonary Nodule  
Dimension: 0.8cm

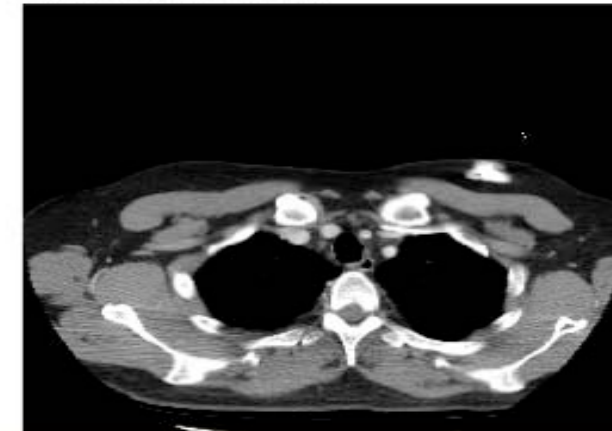


Date: 10/10/06

CT Chest, Abdomen and Pelvis



Pulmonary Nodule  
Dimension: 1.0cm

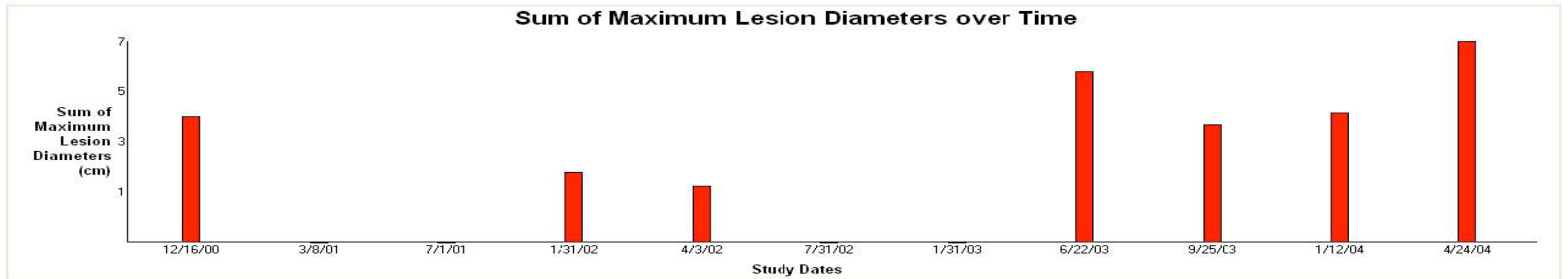
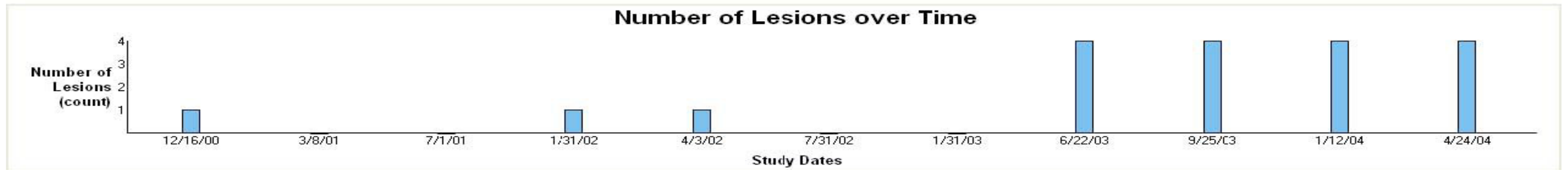


# Levy 2007 - LesionViewer

LesionViewer [minimize] [maximize] [close]

[View Scan Images](#)   
 [View Disease Summary](#)   
 [View Raw Data](#)   
 [Select New Patient](#)

Name: Williams, Laura  
MR Number: 44444444



Study Date	12/16/00	3/8/01	7/1/01	1/31/02	4/3/02	7/31/02	1/31/03	6/22/03	9/25/03	1/12/04	4/24/04
Diagnosis	[remission]	[no disease]	[no disease]	[progression]	[stable]	[remission]	[no disease]	[regression]	[stable]	[progression]	[progression]
Change				+1 lesions	19% shrinkage			+4 lesions	31% shrinkage	9% growth	71% growth

# Earlier Work by Others

- AVT 2009 – Algorithm Validation Toolkit
  - NCI caBIG *in vivo* Imaging Workspace project
  - Measurement Variability Toolkit (MVT) component
    - tabulation and charting
    - interface with R statistics package
  - only supports proprietary NCI AIM format

*Disclosure: author was involved in AVT RFP and sub-contractor to Siemens Corporate Research on AVT use-case development and testing*

<https://wiki.nci.nih.gov/display/AVT/Algorithm+Validation+Toolkit+%28AVT+%29+Project+Pages>



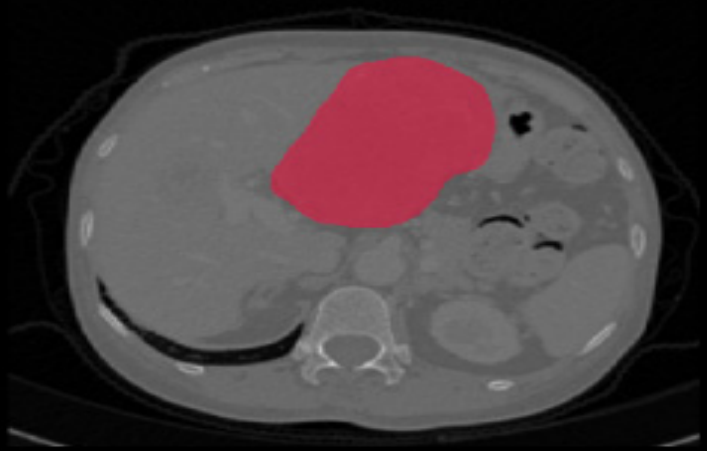
Computation Results

Subject Name	Rel VolDifferen...	SurfDistance(Avera...	Surface Distance(RMS)[...	Surface Distance(Maxim...	Volume Overlap[%]
LTS_IMG01	1.33747	0.33584	0.517345	2	93.38901
LTS_IMG02	78.8215	7.56008	8.99904	18.2481	9.685699
LTS_IMG03	11.4208	0.560206	0.852003	2.6543	70.1677
LTS_IMG04	5.70567	1.76527	2.77575	13.3707	77.781296
LTS_IMG05	24.5516	2.60058	3.84065	13.1668	67.9256
LTS_IMG06	8.13259	0.592667	0.894234	3.15362	78.9365
LTS_IMG07	48.2725	8.57166	10.012	26.0819	47.2082
LTS_IMG08	44.4414	7.37036	8.67253	22.3928	47.052
LTS_IMG09	97.8516	2.70283	3.91401	13.2583	49.0714
LTS_IMG10	25.8335	1.66703	2.21645	7.30861	69.5749
Mean	34.6369	3.37275	4.2694	12.1635	61.0792
SD	32.6547	3.19619	3.63071	8.41635	23.6128
CV	0.942774	0.94765	0.850403	0.691934	0.386592

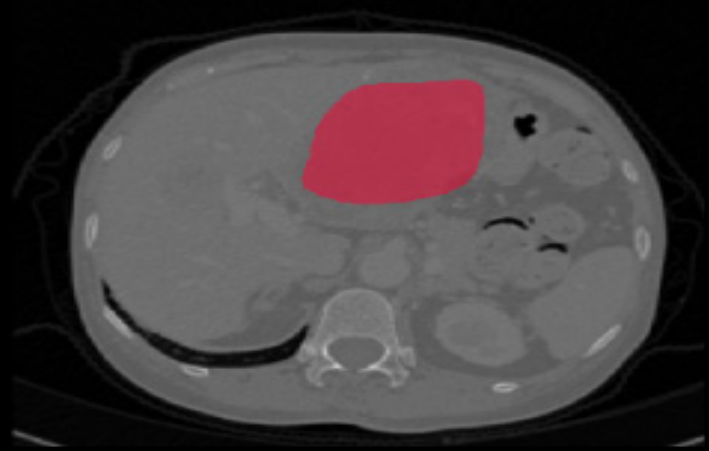
Outliers Plotting

Relative VolumeDifference: Top 25%	Surface Distance(RMS): Top 25%	Volume Overlap: Bottom 25%
LTS_IMG02	LTS_IMG02	LTS_IMG02
LTS_IMG07	LTS_IMG07	LTS_IMG07
LTS_IMG09	LTS_IMG08	LTS_IMG08

Norminal GT



Annotation



# AVT

Statistics Analysis

Methods:

Comparison to be Analyzed	Statistical Method
All Measurements	Mean
All Measurements	SD
All Measurements	CV

Outlier Analysis

Threshold:

Comparison to be Analy...	Outlier Criteria
Relative VolumeDifference	Top 25%
Surface Distance(RMS)	Top 25%
Volume Overlap	Bottom 25%

Plotting

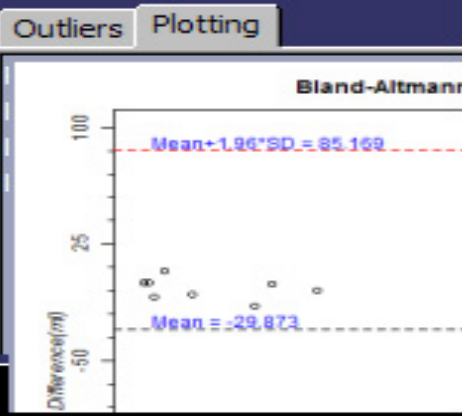
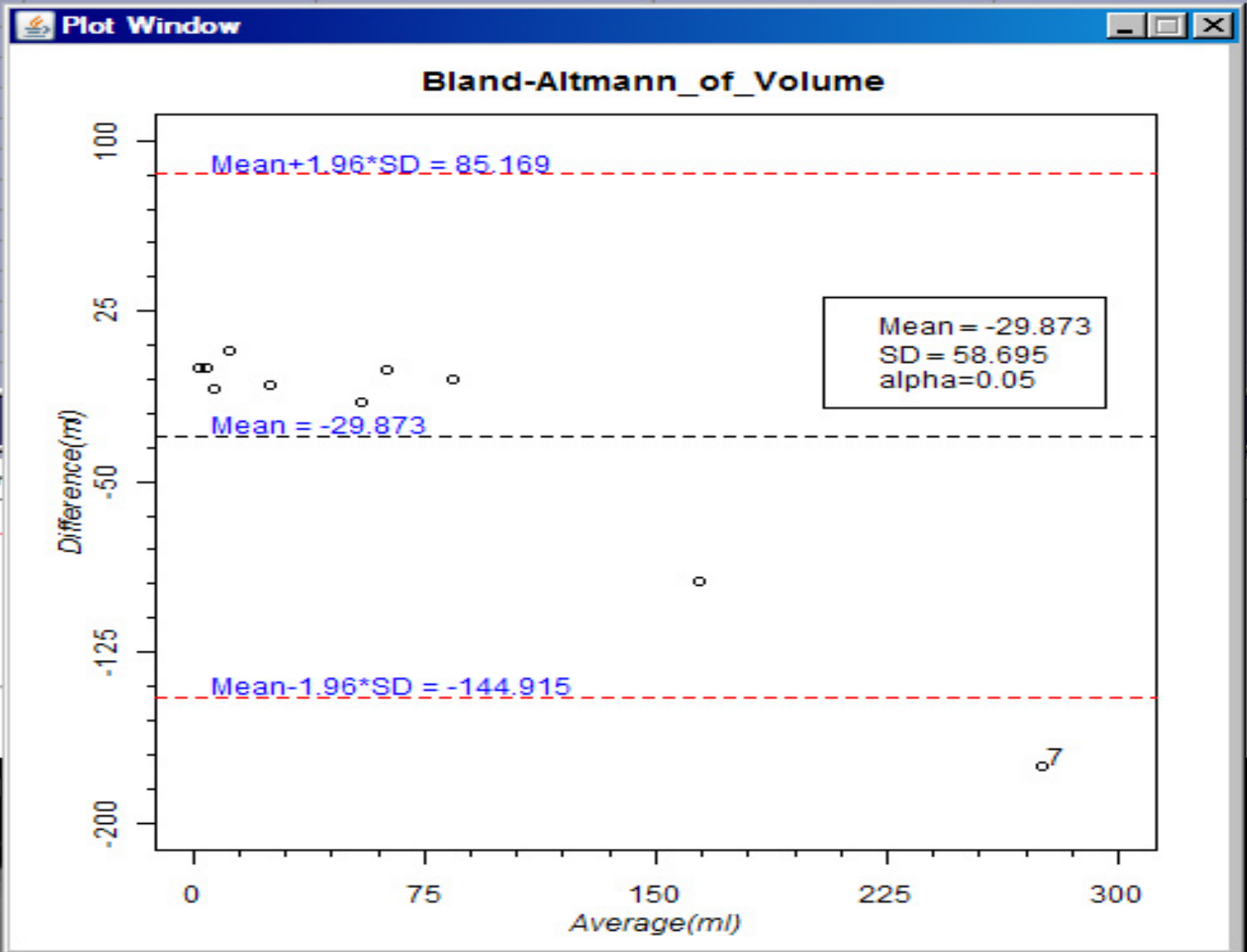
Charts:

Plotting title	Chart
Bland-Altman of Volume	Bland-Altman
Scatter of Volume	Scatter
Surface Distance (RMS)	Scatter

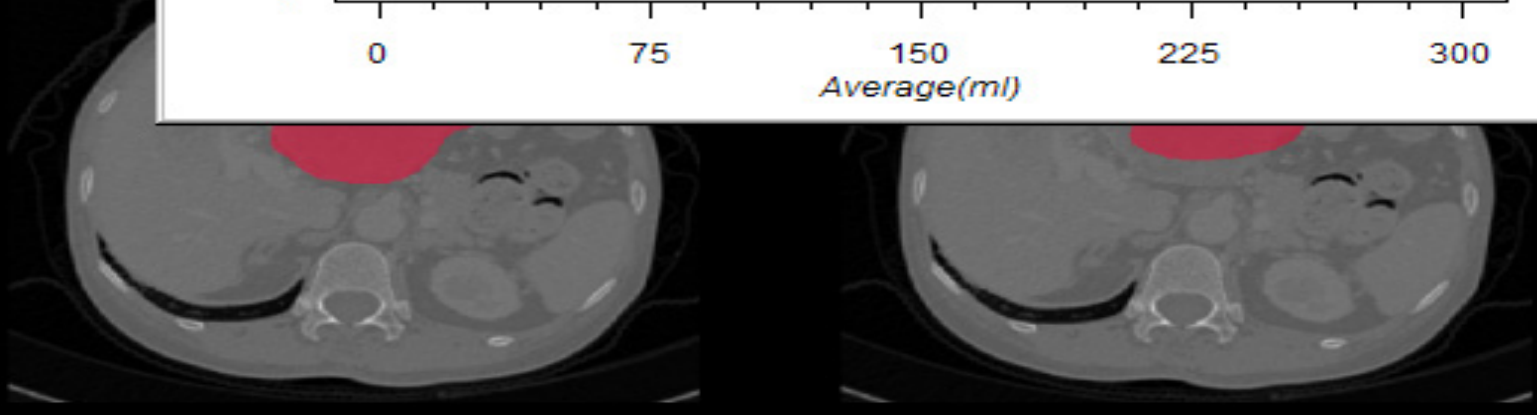
Finish calculation

Computation Results

Subject Name	Rel VolDifferen...	SurfDistance(Avera...	Surface Distance(RMS)[...	Surface Distance(Maxim...	Volume Overlap[%]
LTS_IMG01	1.33747	0.33584	0.517345	2	93.38901
LTS_IMG02	78.8215				
LTS_IMG03	11.4208				
LTS_IMG04	5.70567				
LTS_IMG05	24.5516				
LTS_IMG06	8.13259				
LTS_IMG07	48.2725				
LTS_IMG08	44.4414				
LTS_IMG09	97.8516				
LTS_IMG10	25.8335				
Mean	34.6369				
SD	32.6547				
CV	0.942774				



Normal



# AVT

Statistics Analysis

Methods: Add Custom Del

Comparison to be Analyzed	Statistical Method
All Measurements	Mean
All Measurements	SD
All Measurements	CV

Outlier Analysis

Threshold: Add Del

Comparison to be Analy...	Outlier Criteria
Relative VolumeDifference	Top 25%
Surface Distance(RMS)	Top 25%
Volume Overlap	Bottom 25%

Plotting

Charts: Add Del

Plotting title	Chart
Bland-Altman of Volume	Bland-Altman
Scatter of Volume	Scatter
Surface Distance (RMS)	Scatter

PLOT

BACK

RUN

Finish calculation