

# Imaging of Cancer



## Imaging of Cancer:

Subtitle: What actually happens in a Radiology Department?

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Molecular Imaging Program, NCI

# Imaging of Cancer

- Imaging is a key element of:
  - Screening (e.g. lung cancer, breast cancer)
  - Staging (has it spread locally? Metastasized?)
  - Monitoring of treatment (Better or worse?)
  - Recurrence (Has it come back?)
  - Prognosis (What will happen?)

# The Main Imaging Devices

- Computed Tomography (CT)
- Magnetic Resonance Imaging (MRI)
- Ultrasound (US)
- Single Photon Emission Computed Tomography (SPECT)
- Positron Emission Tomography (PET)
- Optical Imaging

# The Main Imaging Devices

## Quiz: Name that Scanner



CT



MRI



US

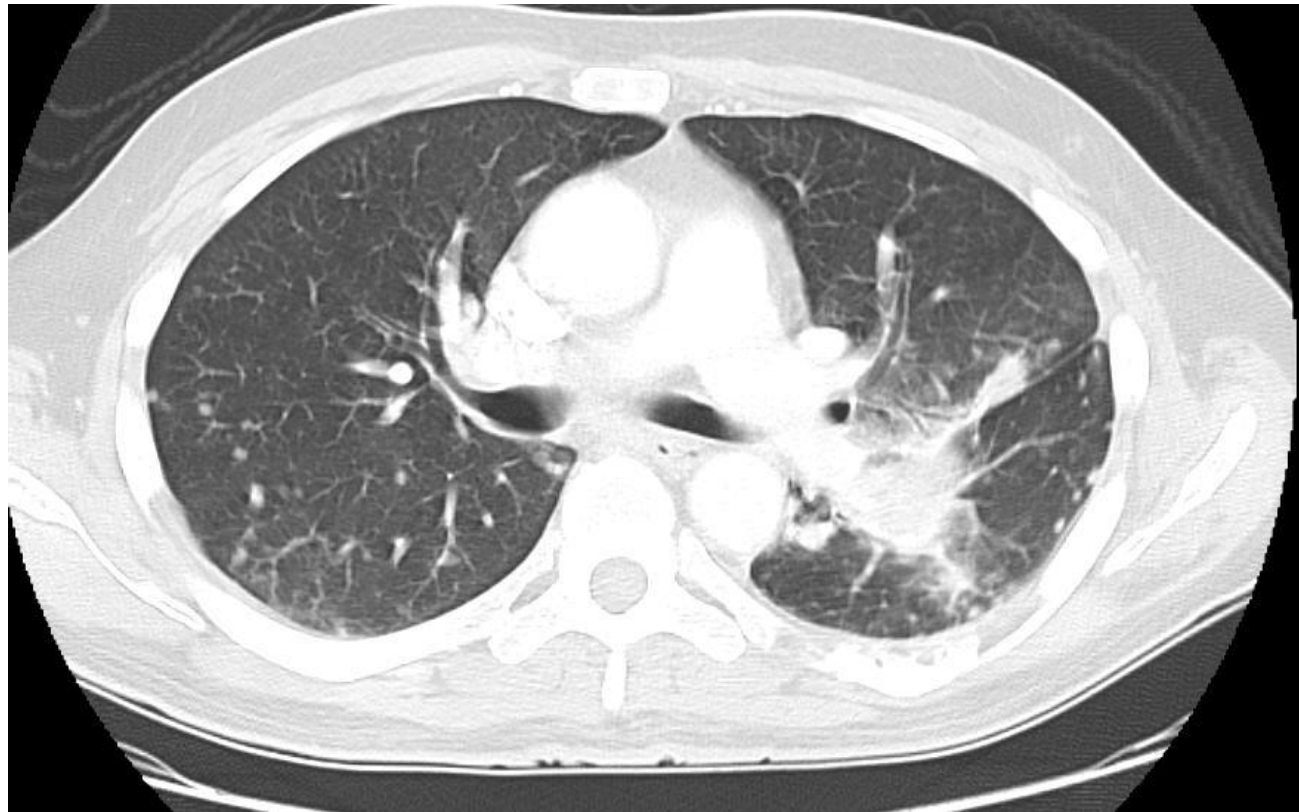


SPECT



PET

# Computed Tomography



# Advantages of CT

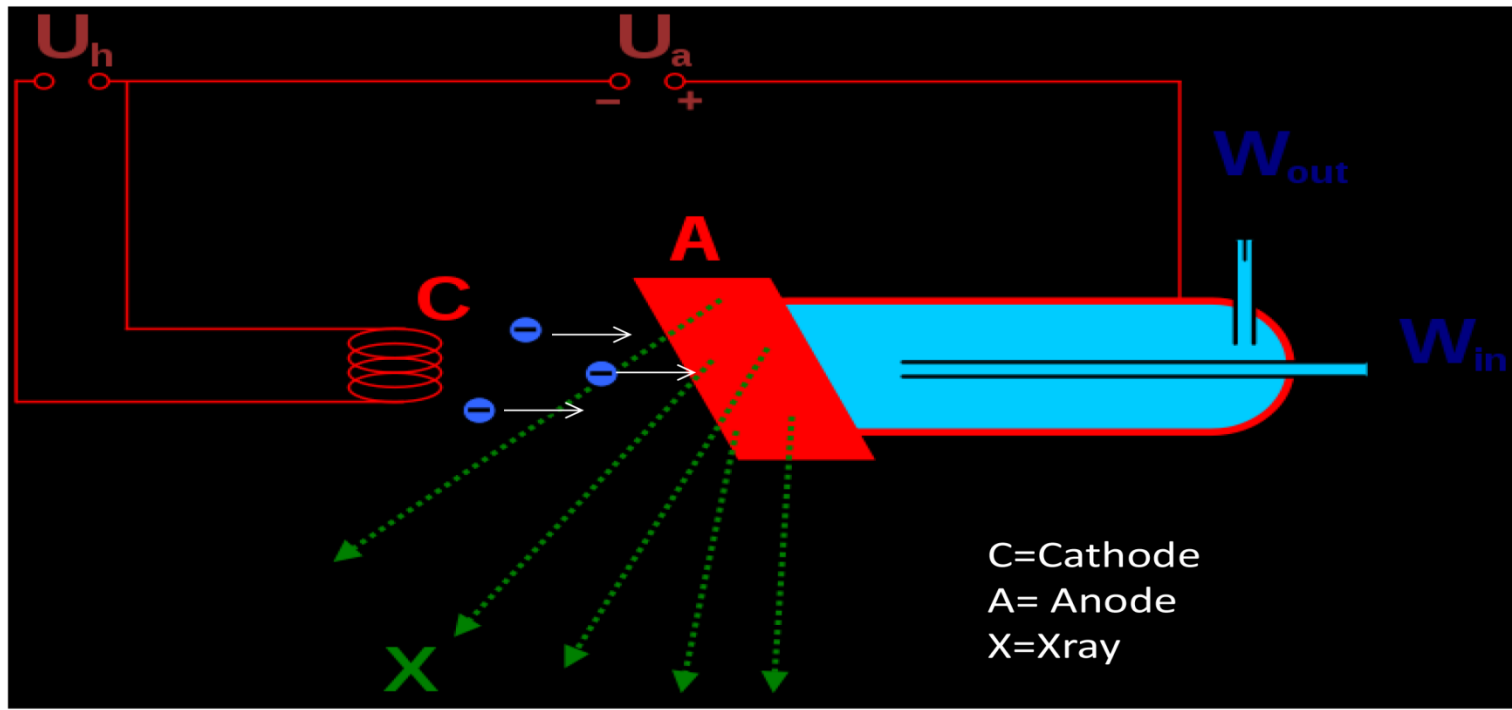
- Widely available
- Minimal prep (NPO, drink contrast)
- Very rapid (2-3 seconds neck to pelvis)
- High resolution
- Relatively inexpensive

# Disadvantages

- **Radiation**
- Often requires iv contrast media
  - Allergic reactions (minimal)
  - Kidney damage (only in high risk patients)
- Anatomic information only

# X-ray production

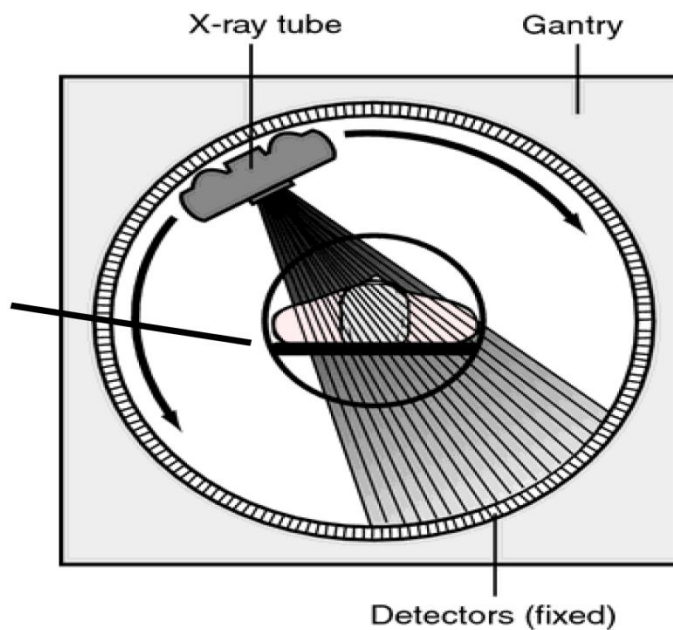
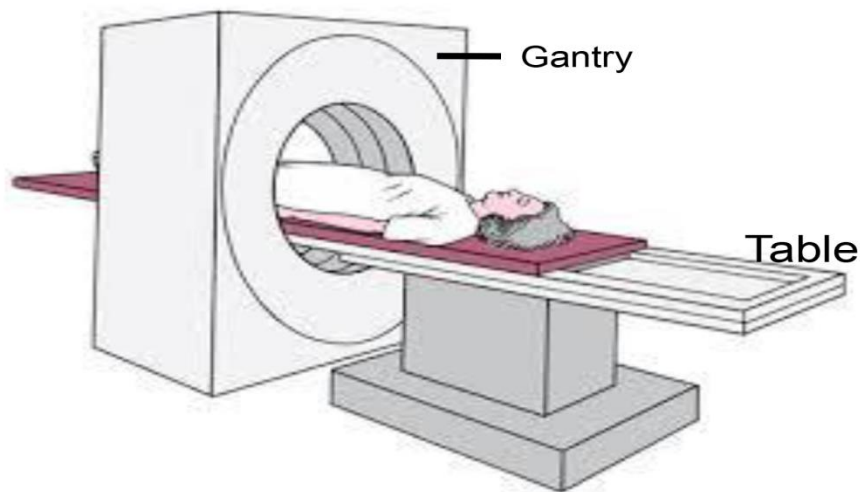
## X-ray production: cathode ray tube





# Basics of CT

# Basics of CT



# CT projection

## Filtered Back Projection

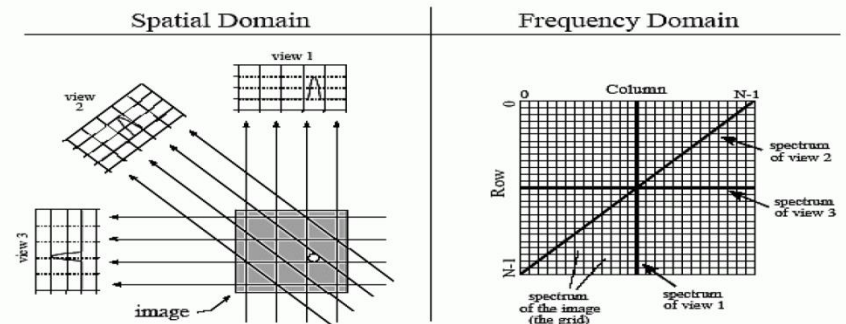
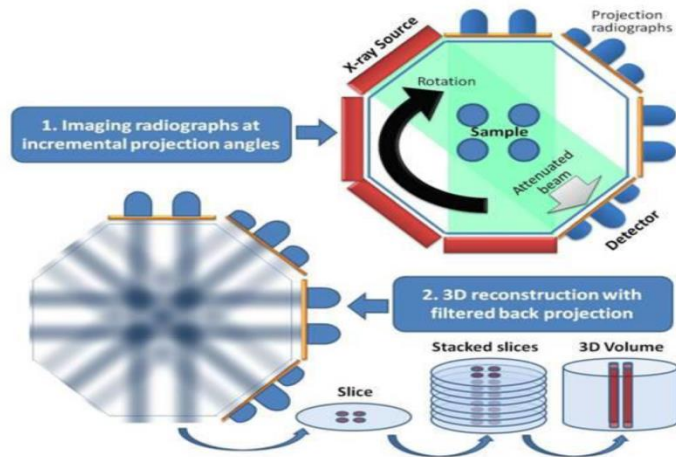
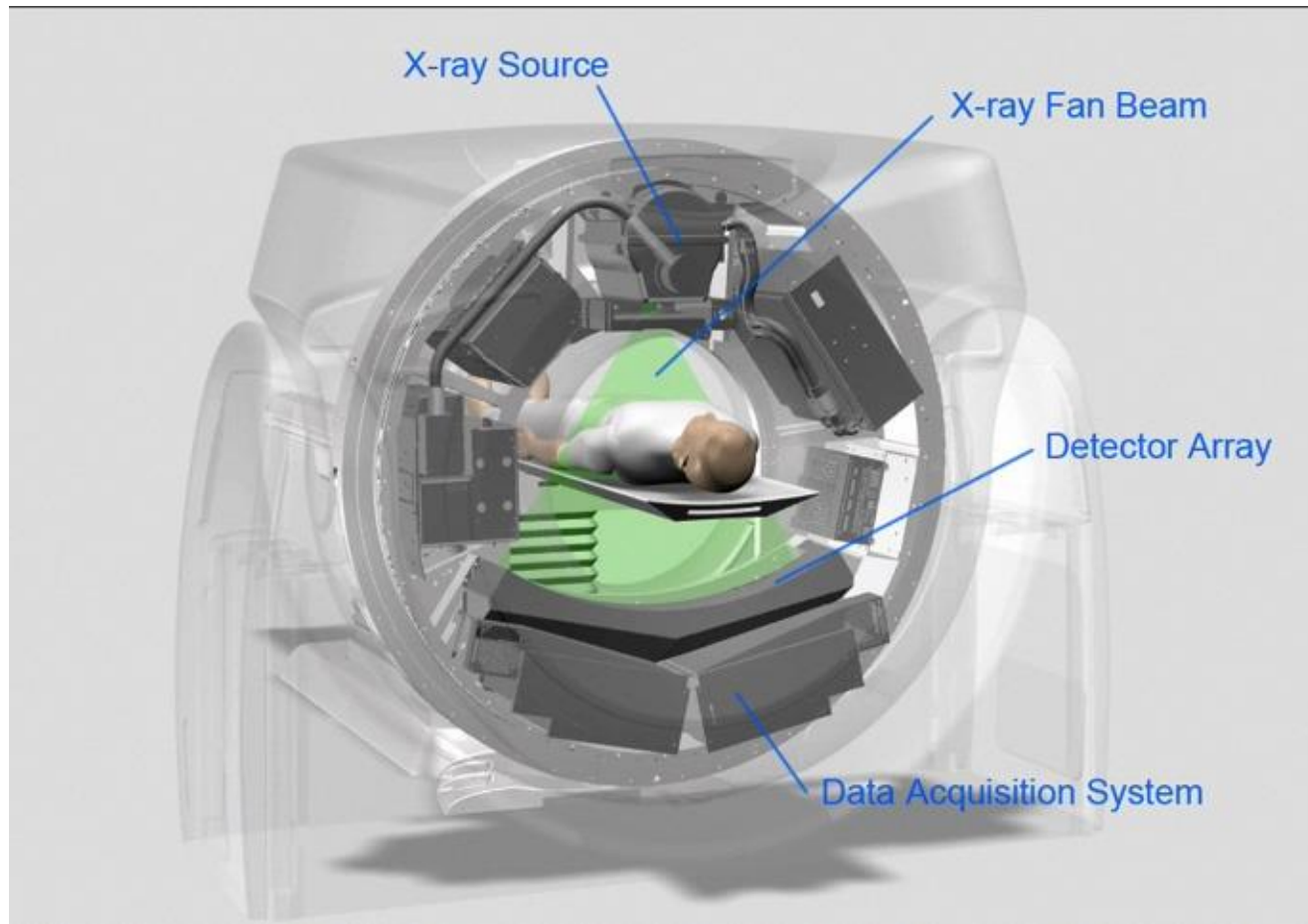
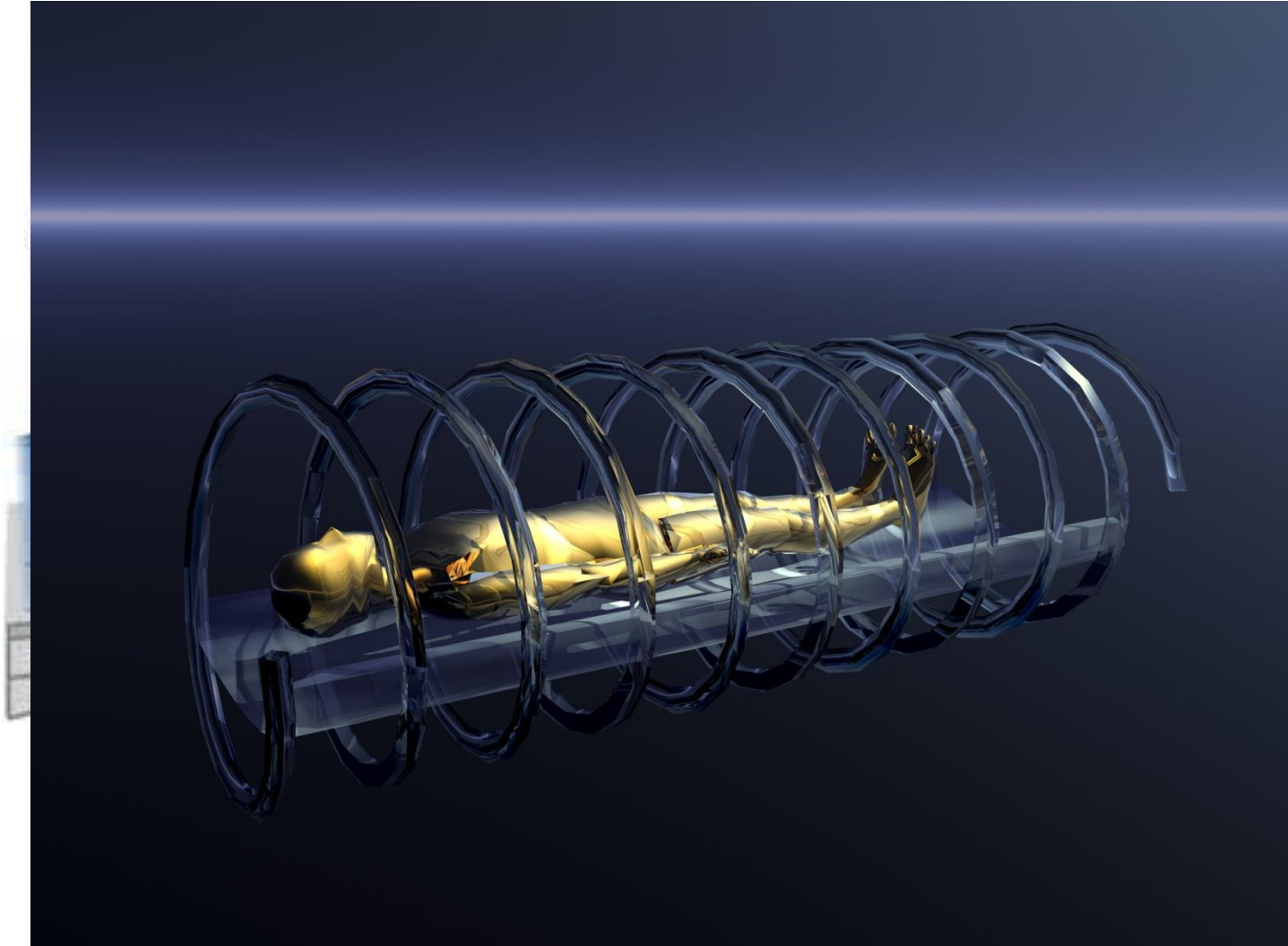


FIGURE 25-18  
The Fourier Slice Theorem. The Fourier Slice Theorem describes the relationship between an image and its views in the frequency domain. In the spatial domain, each view is found by integrating the image along rays at a particular angle. In the frequency domain, the spectrum of each view is a one-dimensional "slice" of the two-dimensional image spectrum.

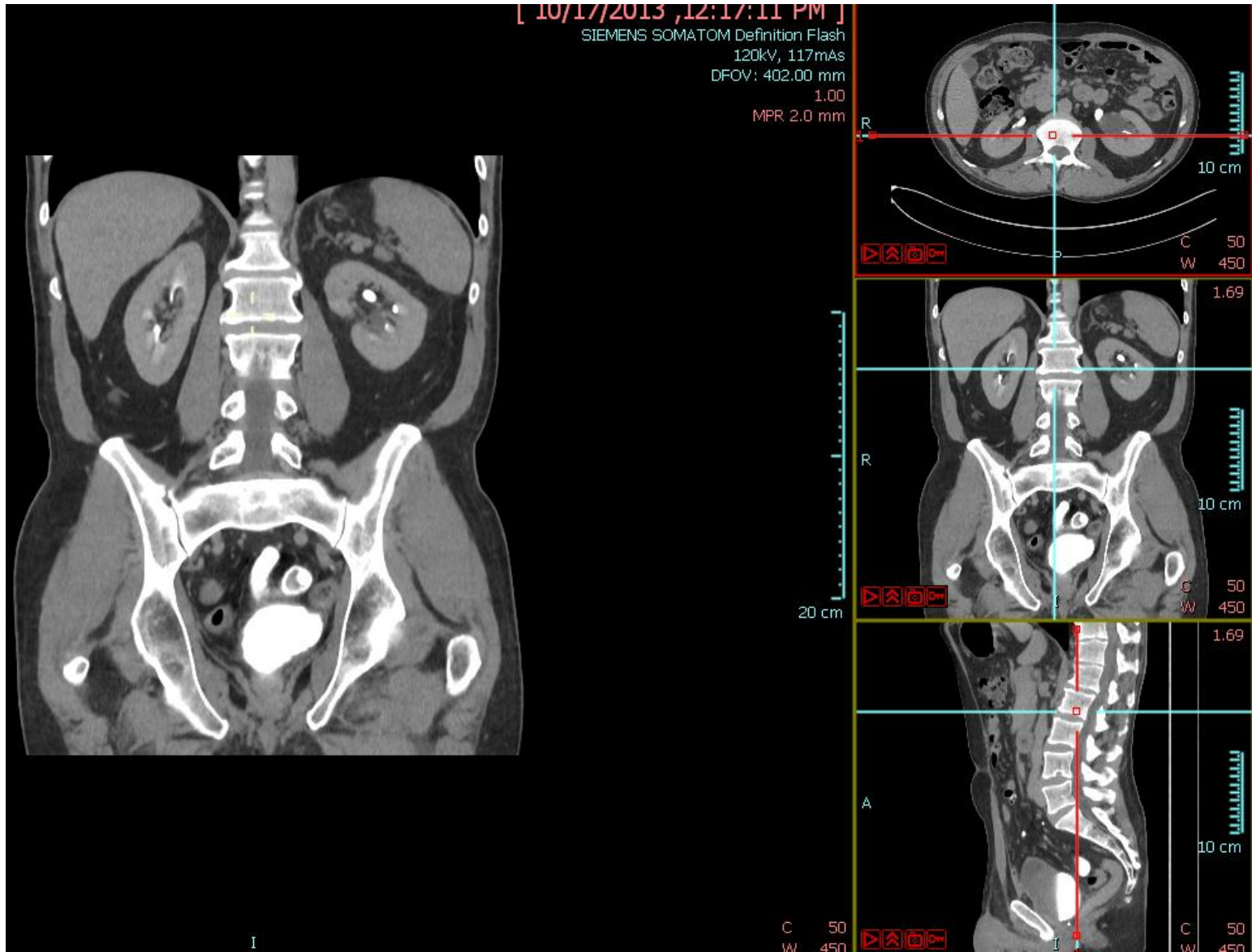
# Cross section of a CT Scanner



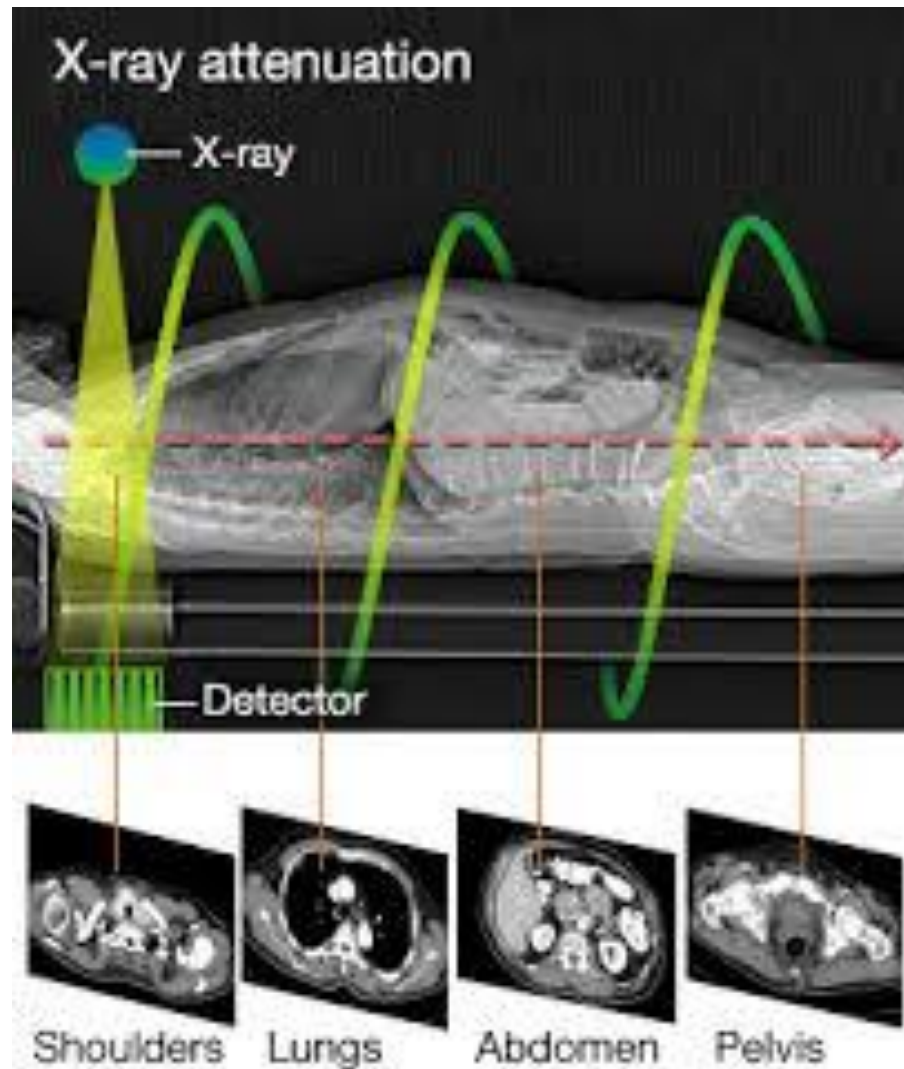
# “Spiral” CT



# “Volume” CT imaging



# Attenuation differences thru the body

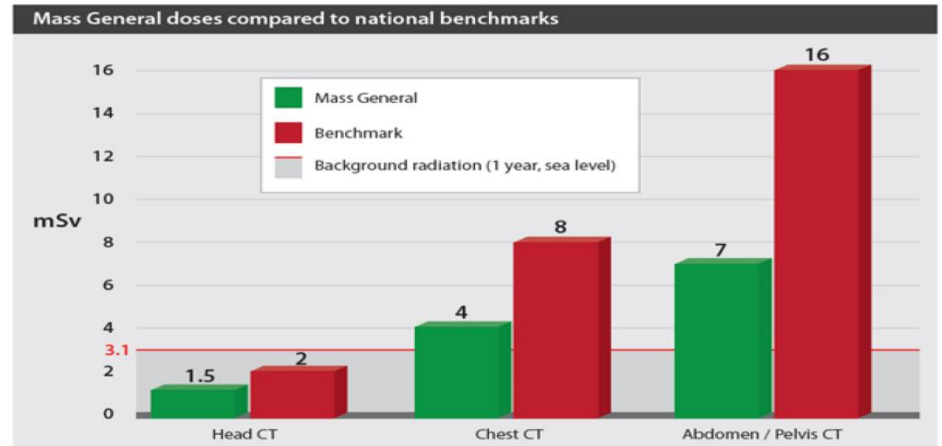


# Radiation

# Radiation

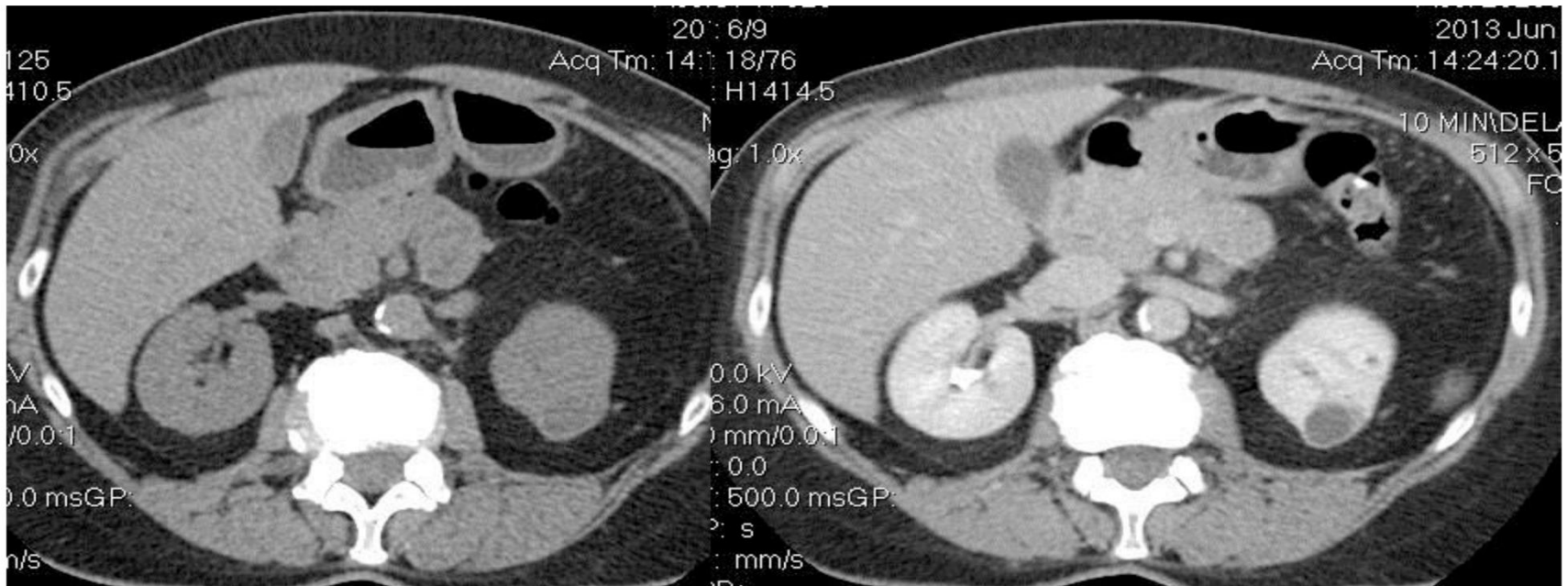


Lower kV (energy) x-rays  
More sensitive detectors  
Better reconstruction algorithms  
“Synthetic” images



# Contrast Media

## Iodinated Contrast Media





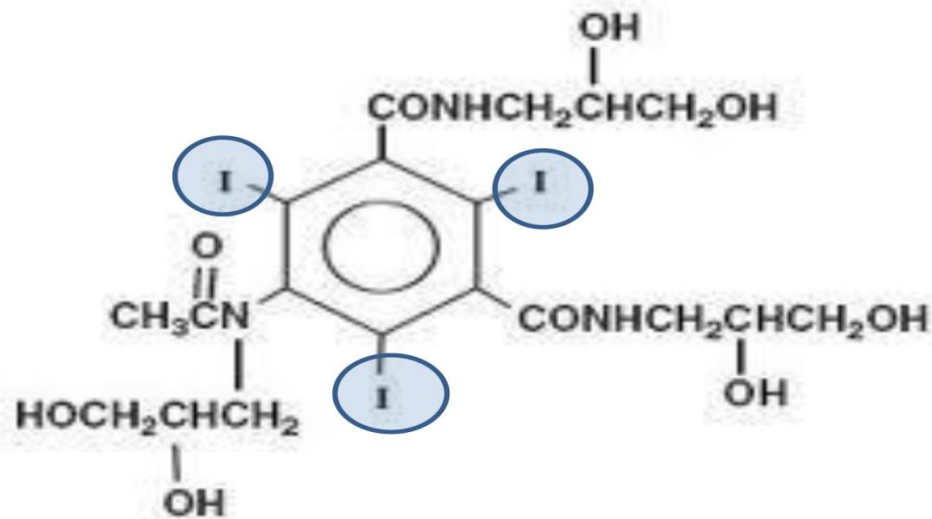
# Iodinated Contrast

# Iodinated Contrast



# Non ionic iodinated contrast

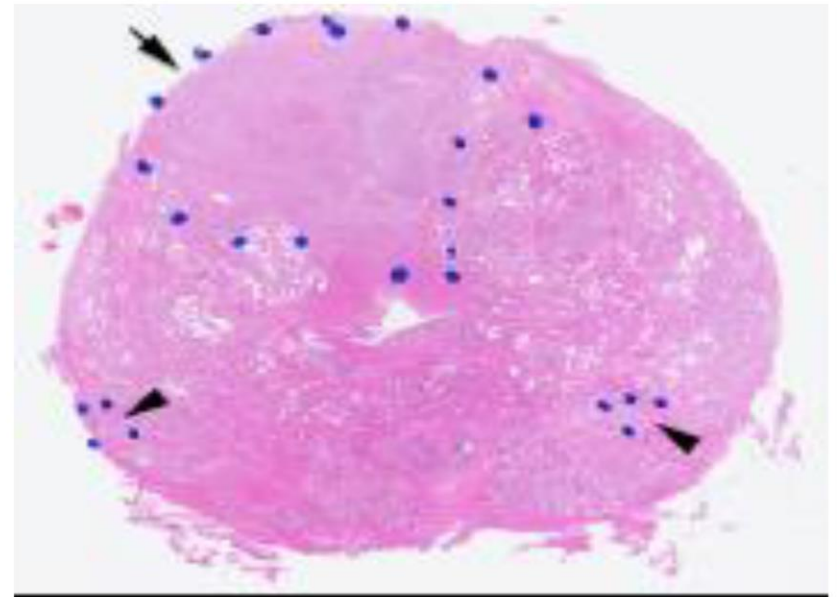
## Non ionic Iodinated Contrast





# MRI

## Magnetic Resonance Imaging



Prostate Cancer on MRI and Pathology

# MRI Advantages

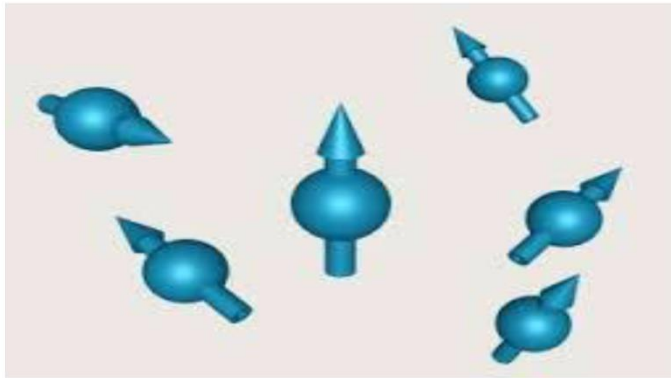
- No radiation
- Multiplanar
- Multiple contrast types:
  - T1 weighting, T2 weighting
  - Diffusion weighting
  - Contrast enhanced MRI
  - Spectroscopy

# MR Disadvantages

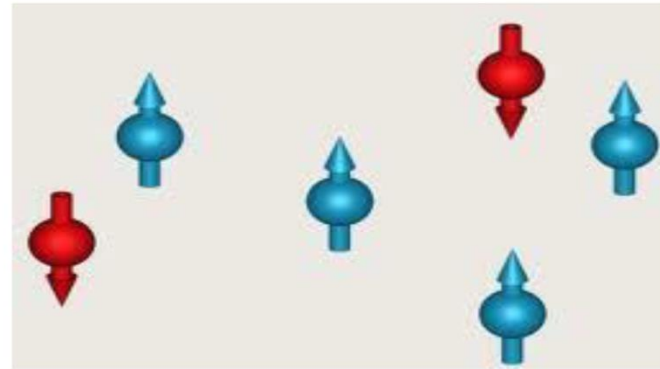
- Slower than CT
- More expensive
- Does not depict calcifications
- Safety issues
  - Metallic objects become projectiles
  - Incompatible with metallic implanted devices
    - Pacemakers
    - Cochlear implants

# MRI physics

## MRI Physics 101



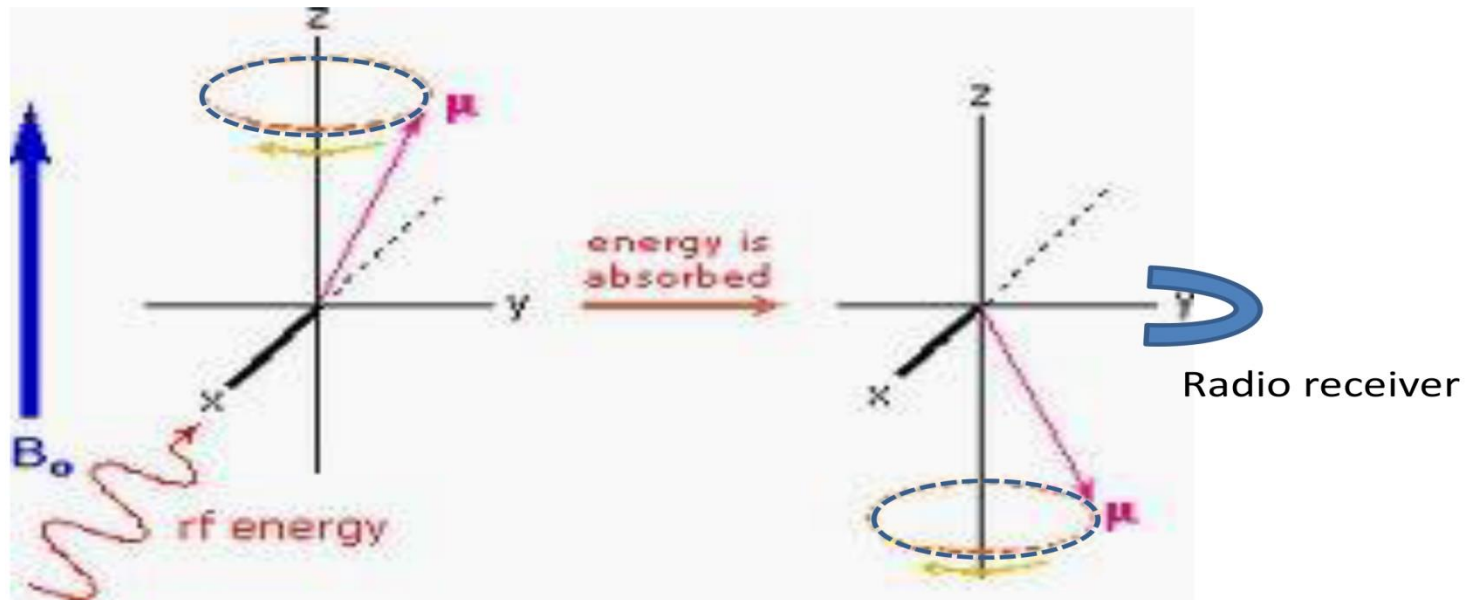
Protons in space: no field



Protons in magnetic field

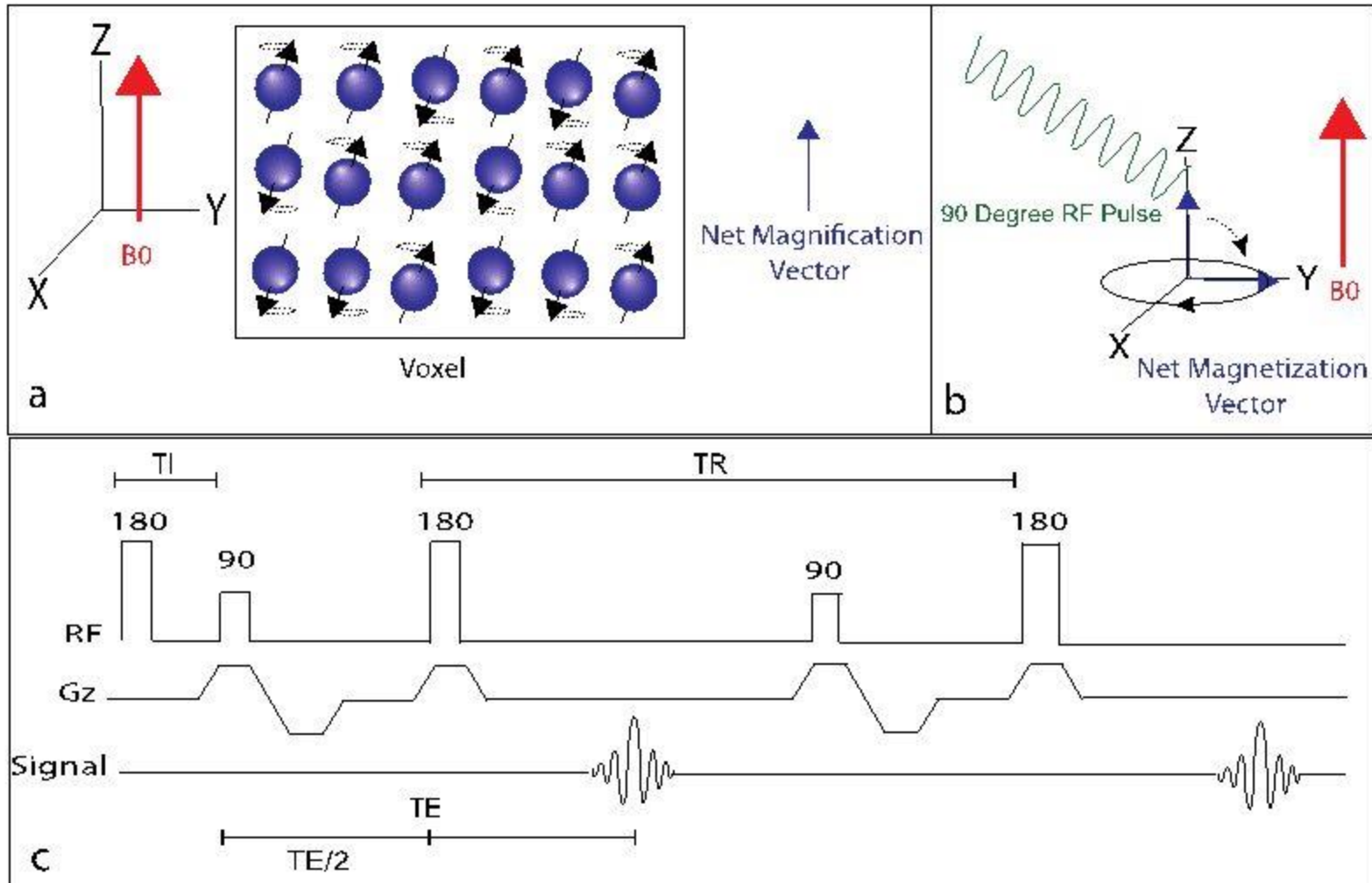
# MR physics

## MR Physics



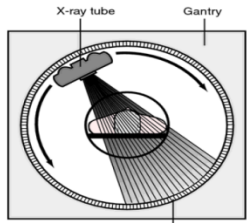


# Summary

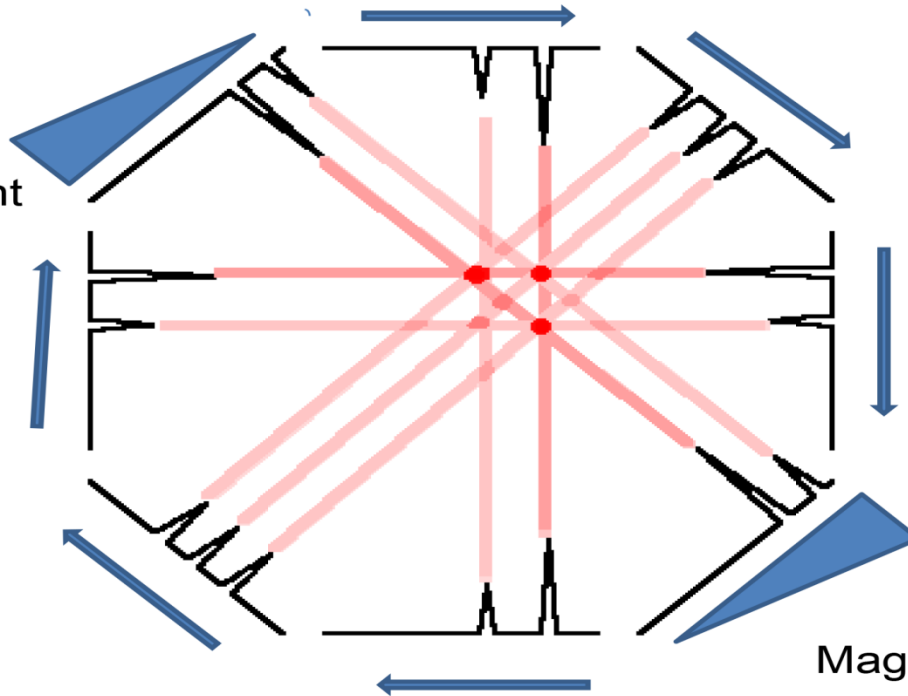


# Creating a MR Image

Creating an MR Image:  
No detectors! Just antennas (coils)

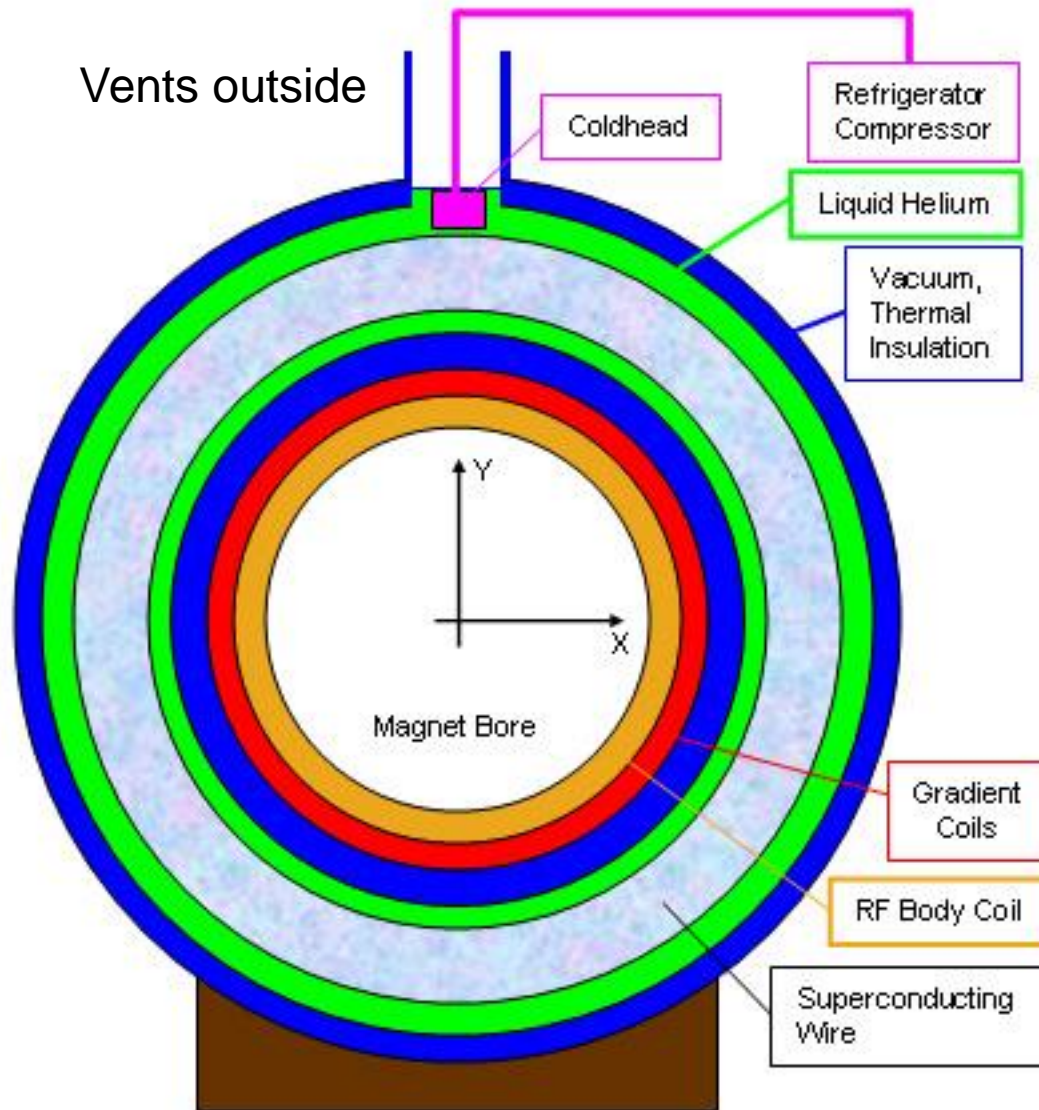


Magnetic Gradient

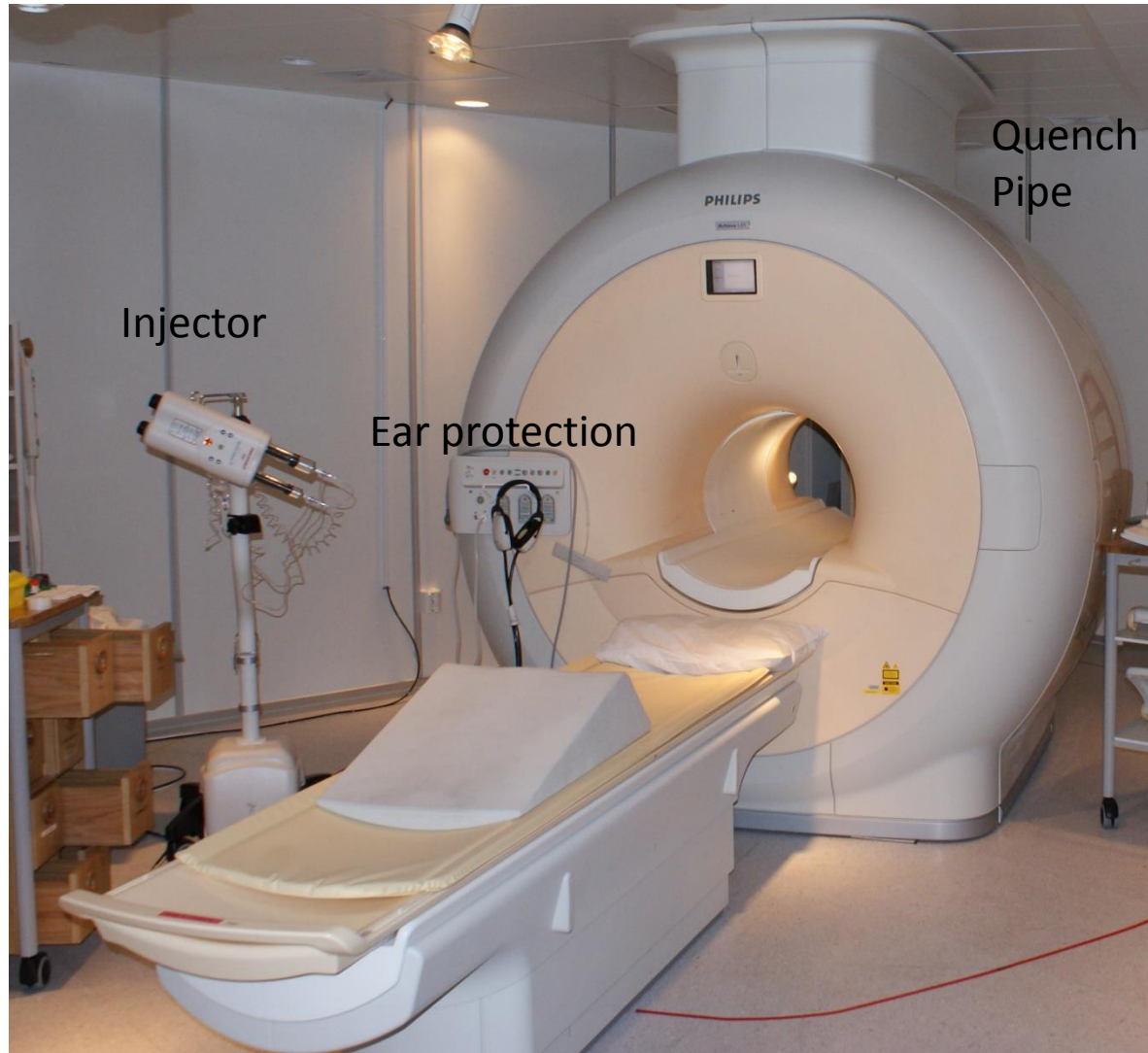


Magnetic Gradient

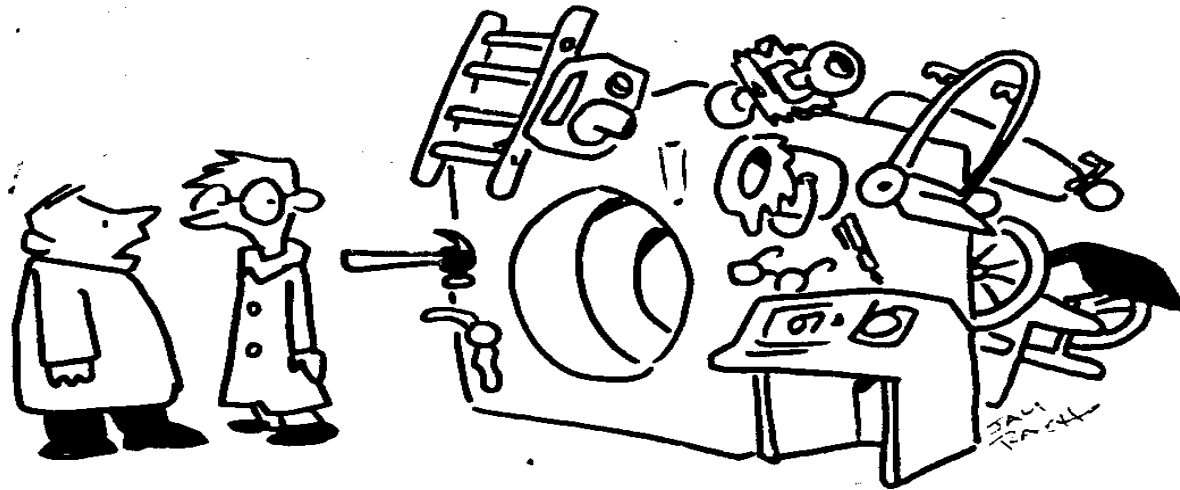
# Anatomy of an MRI



# Safety issues in MRI



# The Importance of MR Safety

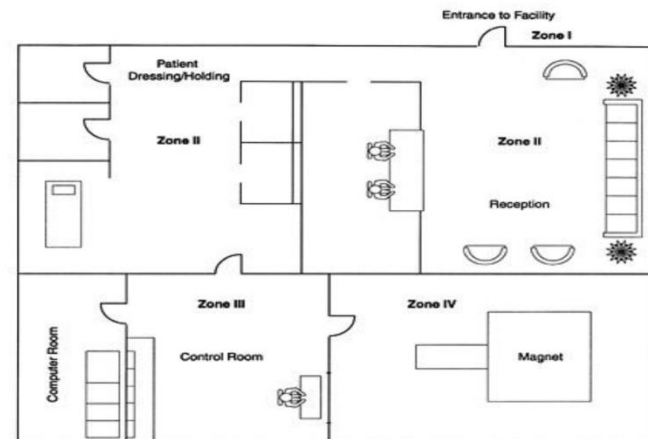


*“Maybe it is time for us to review the magnet safety instructions.”*

# MRI Safety

## MRI SAFETY

- MRI scanners are extremely powerful
- Objects that are attracted by the MRI magnetic field can reach 60 miles per hour.
- A sharp or heavy object can be deadly to anyone standing in its path.
- Metal objects used everyday (scissors, oxygen tanks, infusion pumps, etc) become projectiles
- This can cause potential injury to patients or hospital staff.
- MRI departments are divided into Zones for Safety



# MRI Safety

## MRI SAFETY



# Oxygen tank

## O2 Tank, "Missile"



An Oxygen tank can become an Airborne torpedo in an MRI



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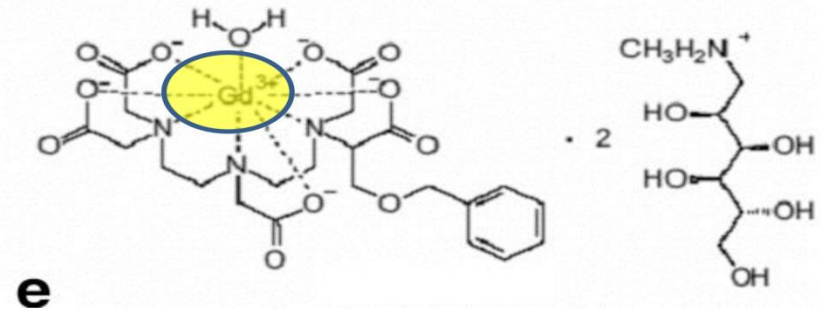
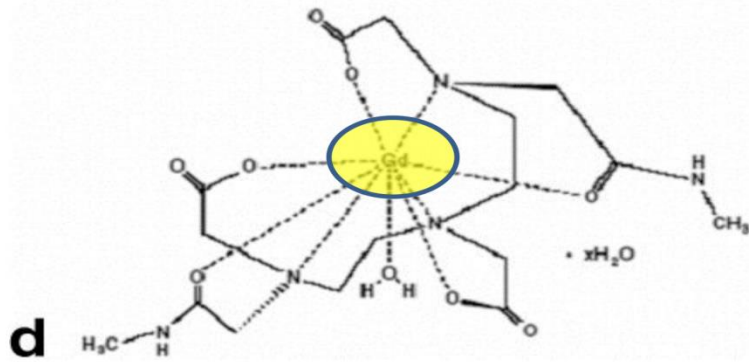
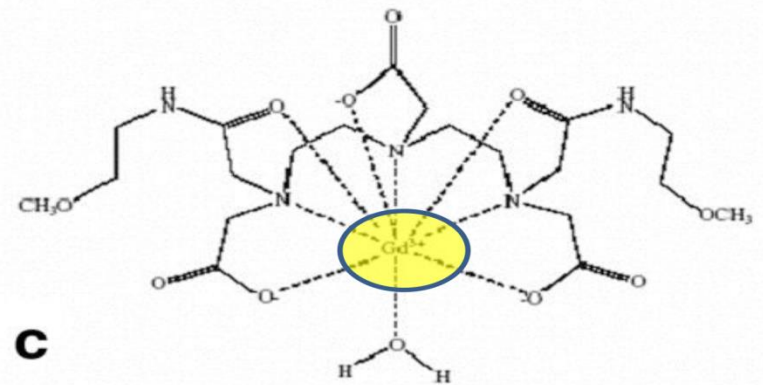
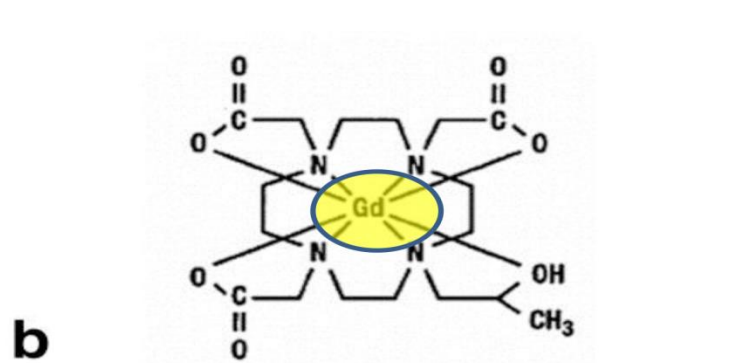
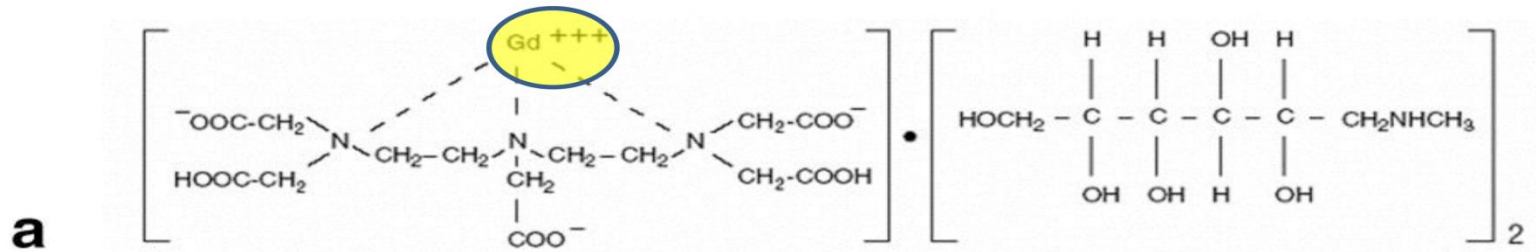
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# Value of Contrast Media



# Gd reagents



# GD reagents

## GD Reagents

Extracellular Gd-CM	Type	Thermodynamic stability constant	Conditional Stability	Amount of excess chelate (mg ml <sup>-1</sup> )	Kinetic stability (dissociation half-life at pH 1.0)
Gadoversetamide, Gd-DTPA-BMEA (OptiMark, Tyco, St. Louis, MO)	Non-ionic linear	16.6	15	28.4	Not available
Gadodiamide, Gd-DTPA-BMA (Omniscan, GE, Waukesha, WI)	Non-ionic linear	16.9	14.9	12	35 s
Gadobutrol, Gd-BT-DO3A (Gadovist, Schering, Berlin, Germany)	Non-ionic cyclic	21.8	Not available	Not available	5 min
Gadoteridol, Gd-HP-DO3A (Prohance, Bracco, Italy)	Non-ionic cyclic	23.8	17.1	0.23	3 h
Gadopentetate Gd-DTPA (Magnavist, Schering, Berlin, Germany)	Ionic linear	22.1	18.1	0.4	10 min
Gadobenate, Gd-BOPTA, (Multihance, Bracco, Italy)	Ionic linear	22.6	18.4	None	Not available
Gadoterate, Gd-DOTA (Dotarem, Guerbet, France)	Ionic cyclic	25.8	18.8	None	> 1 month

# Nephrogenic systemic sclerosis

Examples: nephrogenic systemic sclerosis

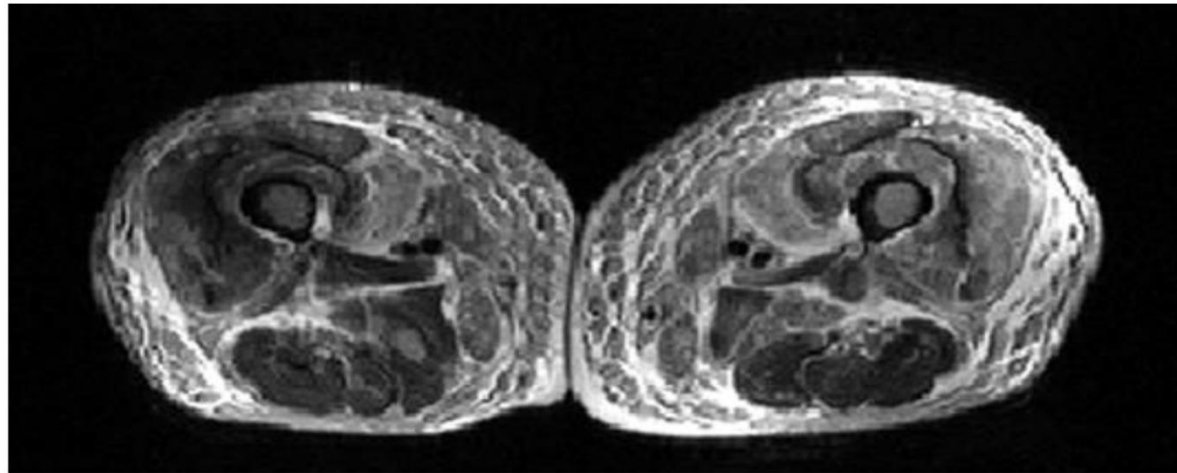
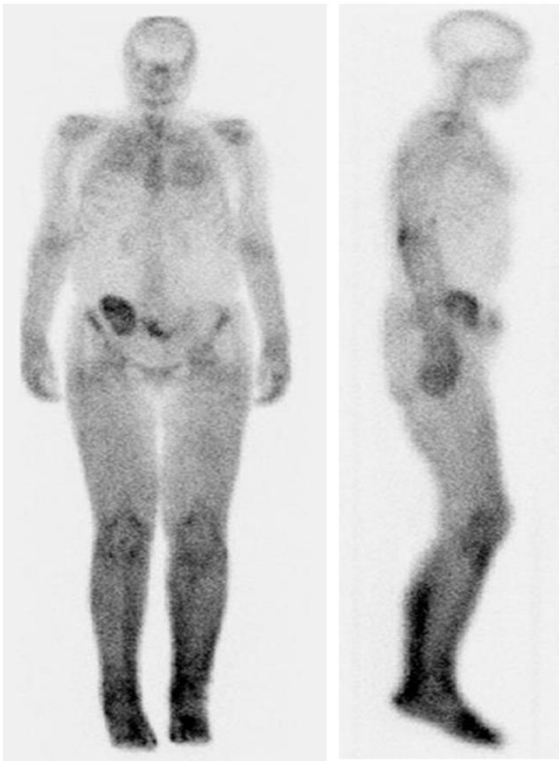


# Nephrogenic Systemic Fibrosis (NSF)

- May 2006 Danish Medicine Agency reported 25 cases of NSF in patients in renal failure who received gadodiamide (~2m)
- Nov 2006 Loma Linda reported 12 (8 on dialysis) NSF patients receiving 0.2mmol/Kg gadodiamide within 2-8 weeks of injection
- Since then over 200 cases have been reported with all of the available contrast agents but mostly Omniscan and Optimark
- By 2013 almost no cases are reported

# Case of NSF

# Case of NSF



# Mechanism

- Gadolinium is highly toxic
- Patients with normal renal function excrete Gd-chelates within 24-48h
- Patients with abnormal renal function may take weeks to excrete the agent
- Dissociation of Gd from the chelate could deposit in soft tissues (documented)
  - Hugh et al. Tissue Gd conc .14-24 ng/mL
- Fibrosis is an inflammatory response to toxic Gd ion.

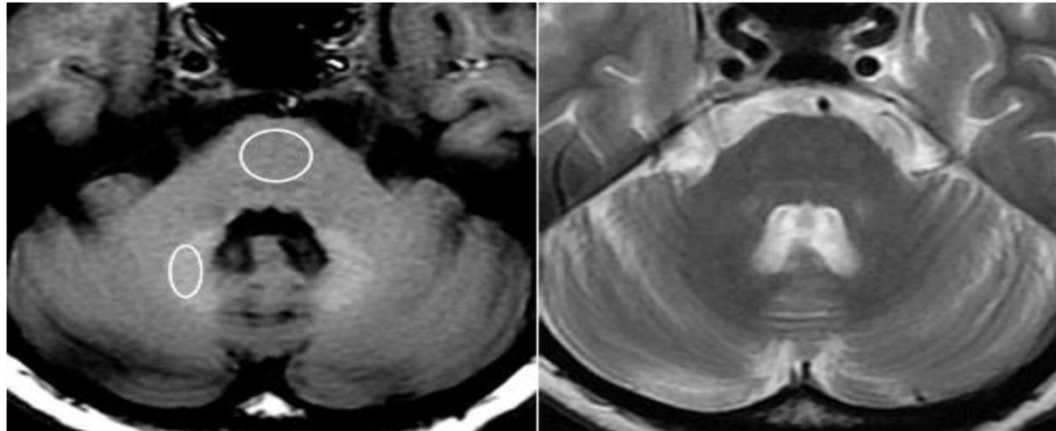
# Risk Factors

- Renal failure:
  - Dialysis
  - Chronic renal failure (GFR <30cc/min)
- Dose
  - Double, triple (vs. half dose)
- Contrast agent
  - Omniscan>Optimark>Magnevist>Prohance>
- Alternative imaging?
  - Non con MRI
  - CT, US, PET



# Residual Gd

## Residual Gadolinium!



Extracellular Gd-CM	Type	Thermodynamic stability constant	Conditional Stability	Amount of excess chelate (mg ml <sup>-1</sup> )	Kinetic stability (dissociation half-life at pH 1.0)
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Gadobenate, Gd-BOPTA, (Multihance, Bracco, Italy)	Ionic linear	22.6	18.4	None	Not available
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# Ultrasound



# US advantages

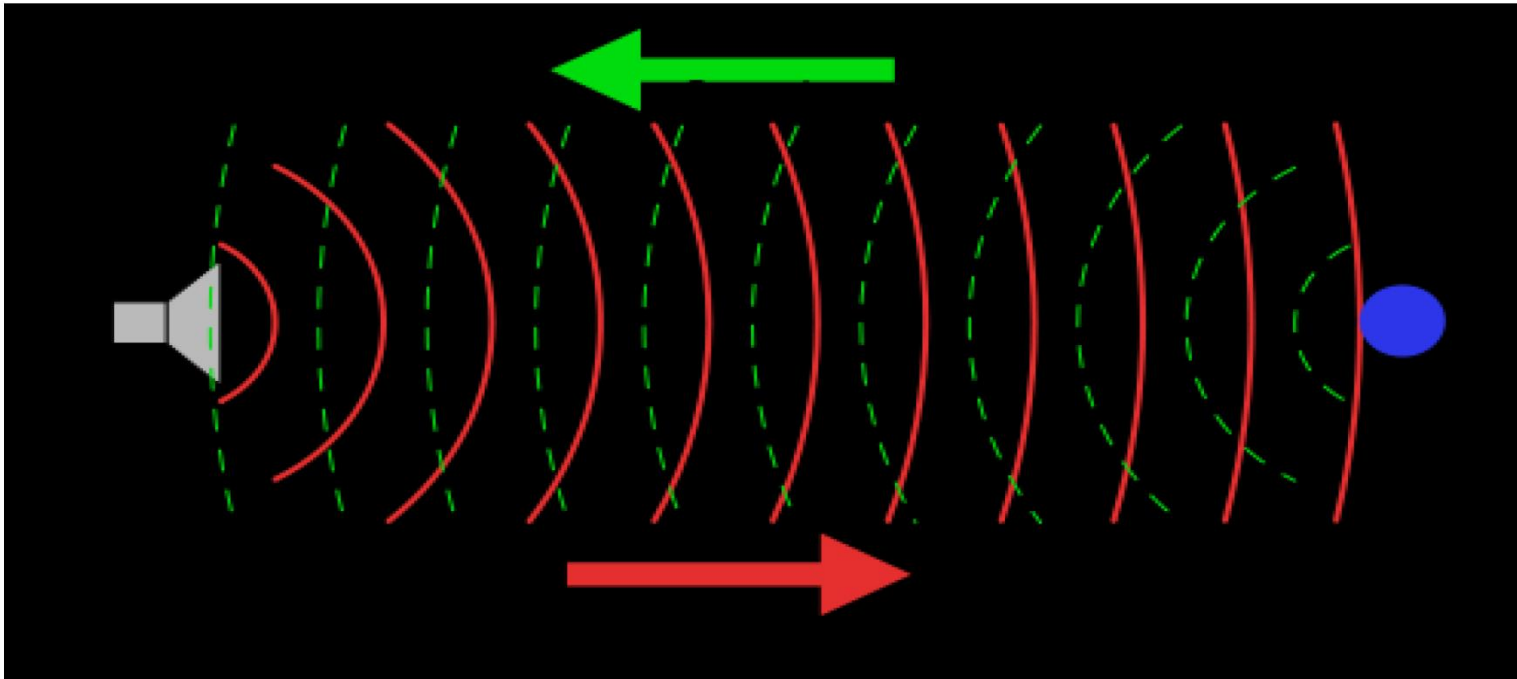
- No radiation
- Real time
- Inexpensive
- Quick, little prep
- No injection

# US disadvantages

- Operator dependent
- What you see is all there is
- Difficult to quantify
- Limited access (lungs, brain, bone etc.)

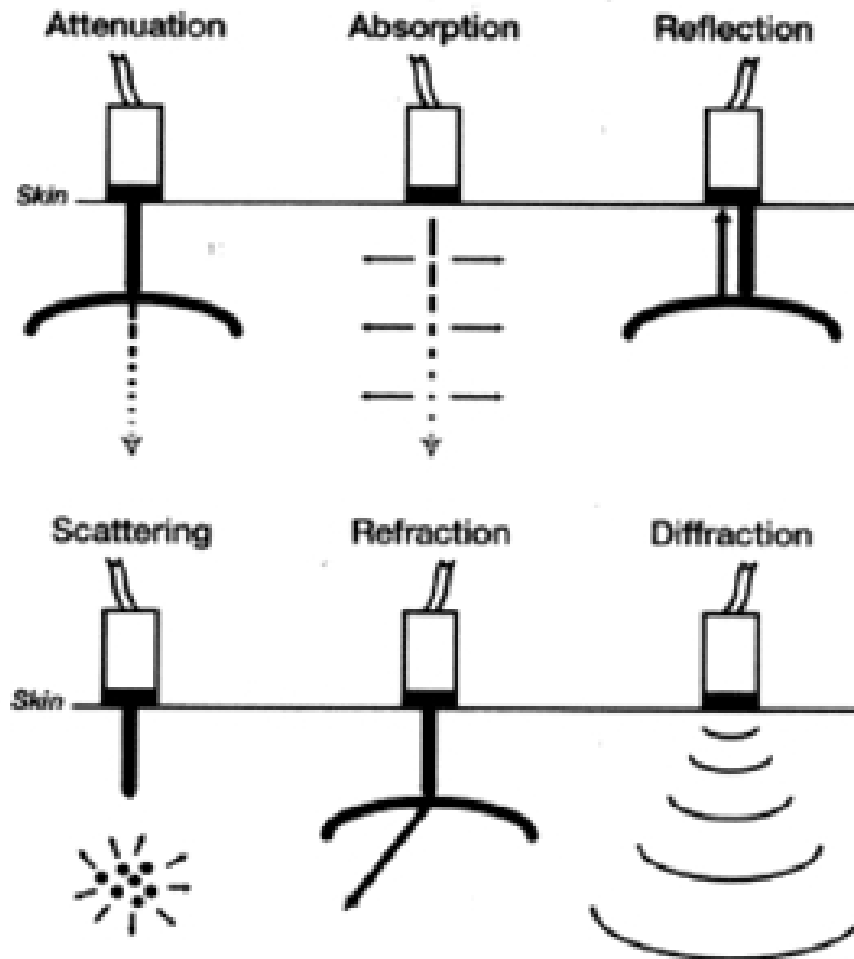
# Ultrasound basics

## US basics

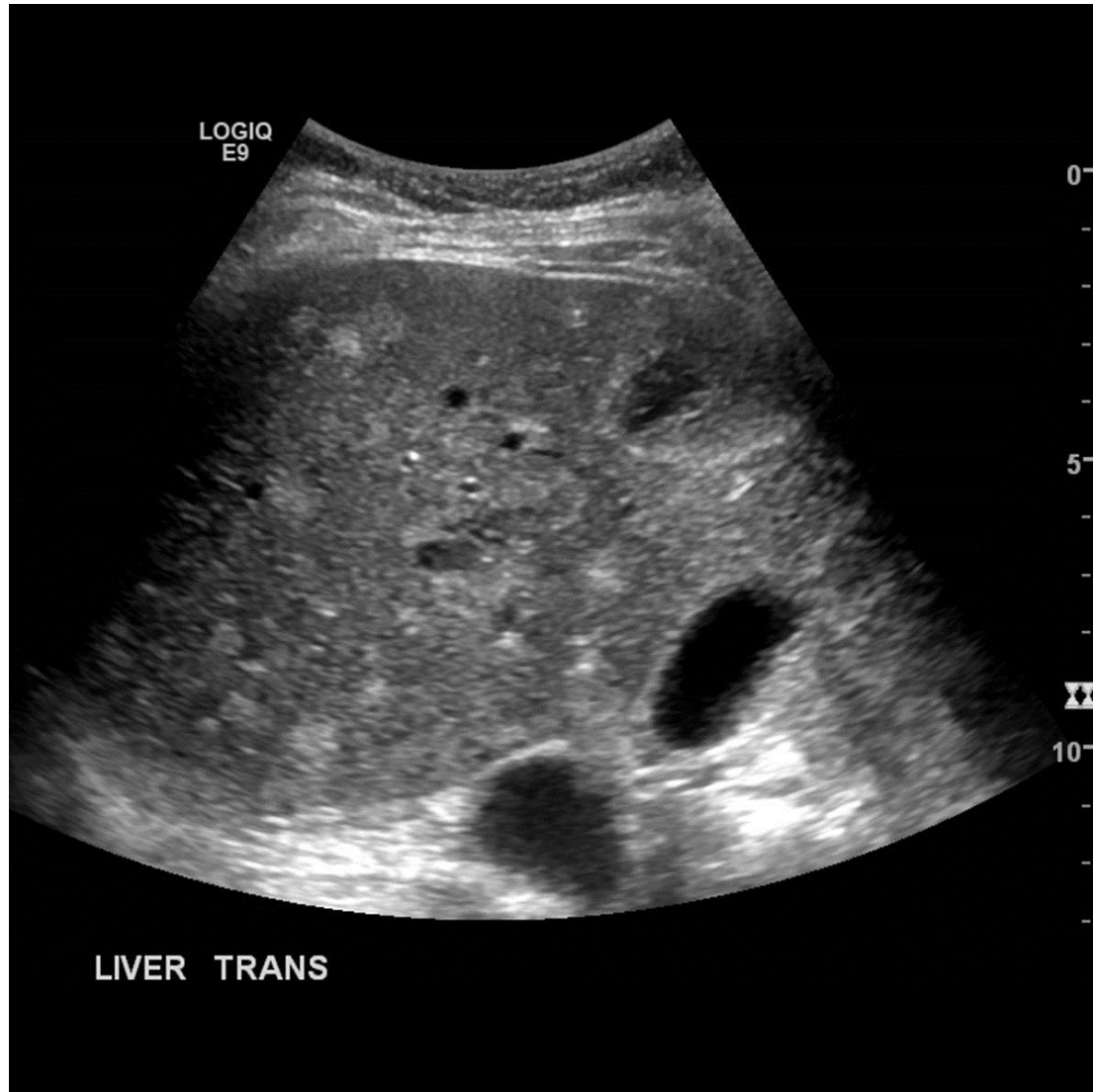


Imaging dependent on the speed of sound  
In tissue

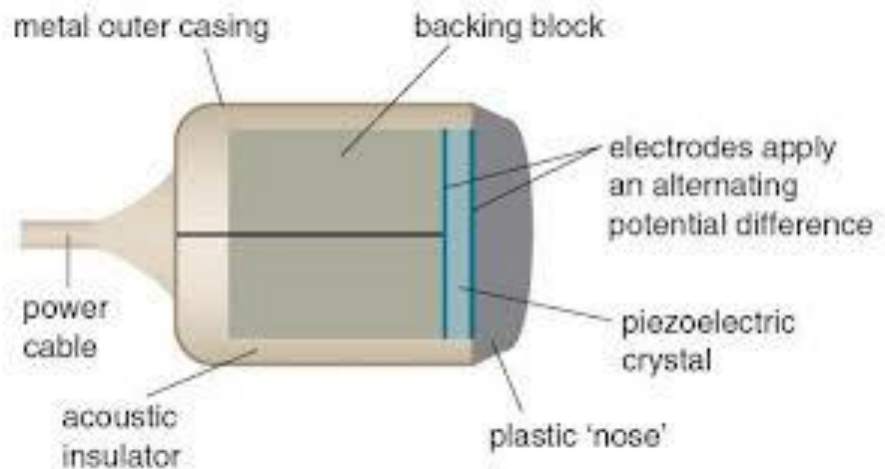
# Fate of sound waves in body



# Liver metastases



# US Probes





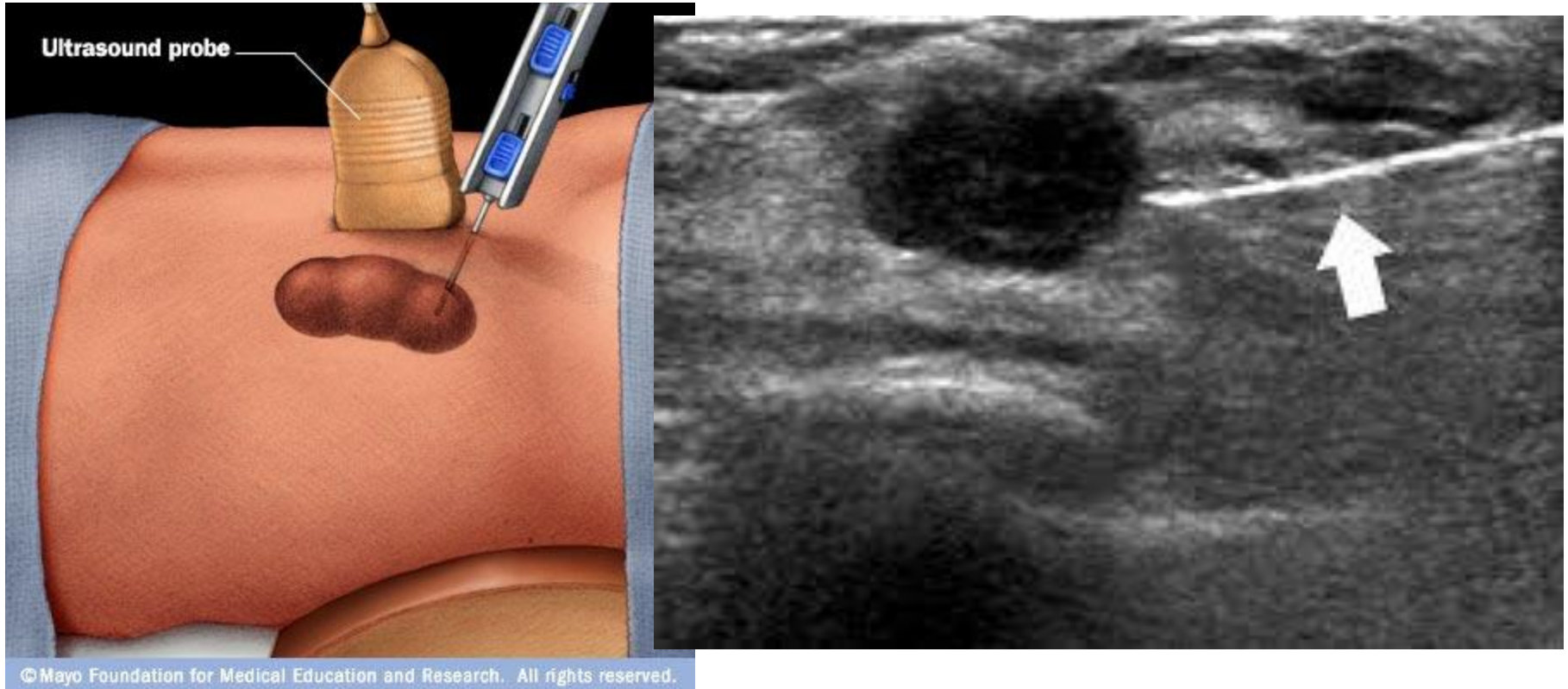


# Evolution of US devices



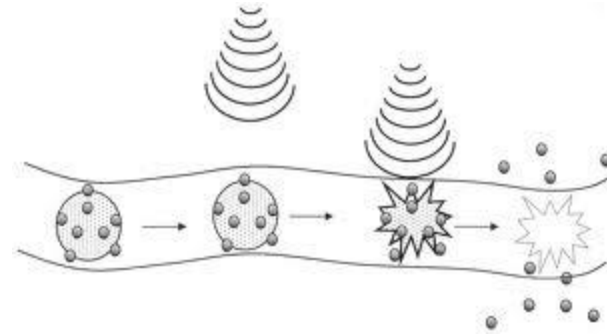
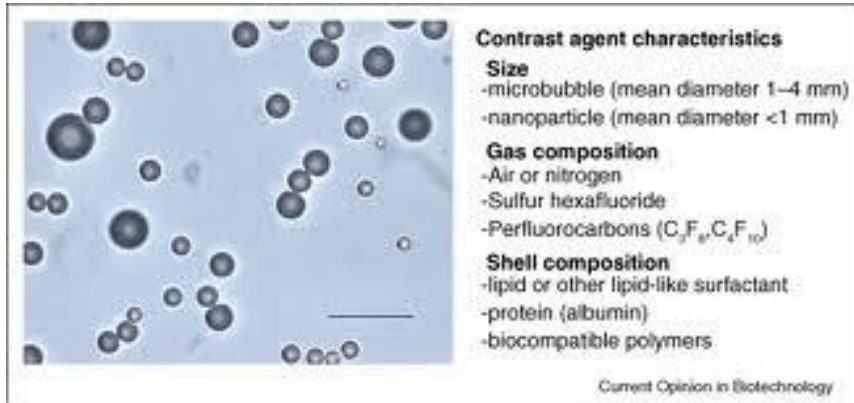


# US guided biopsy-real time





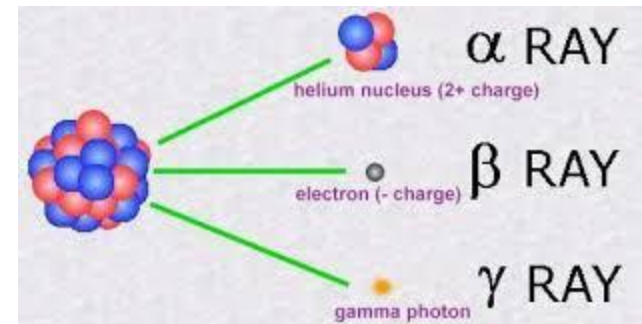
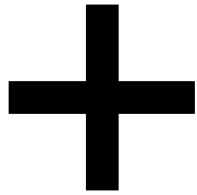
# US Microbubble contrast



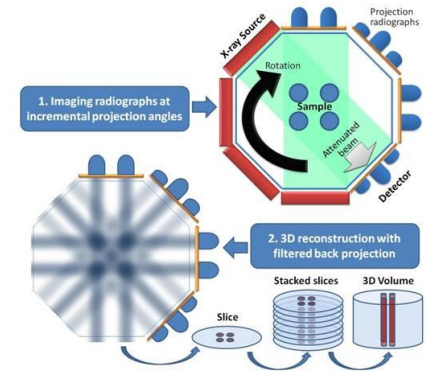


# Single Photon Emission Computed Tomography-SPECT

- Single Photon Emission



- Computed Tomography

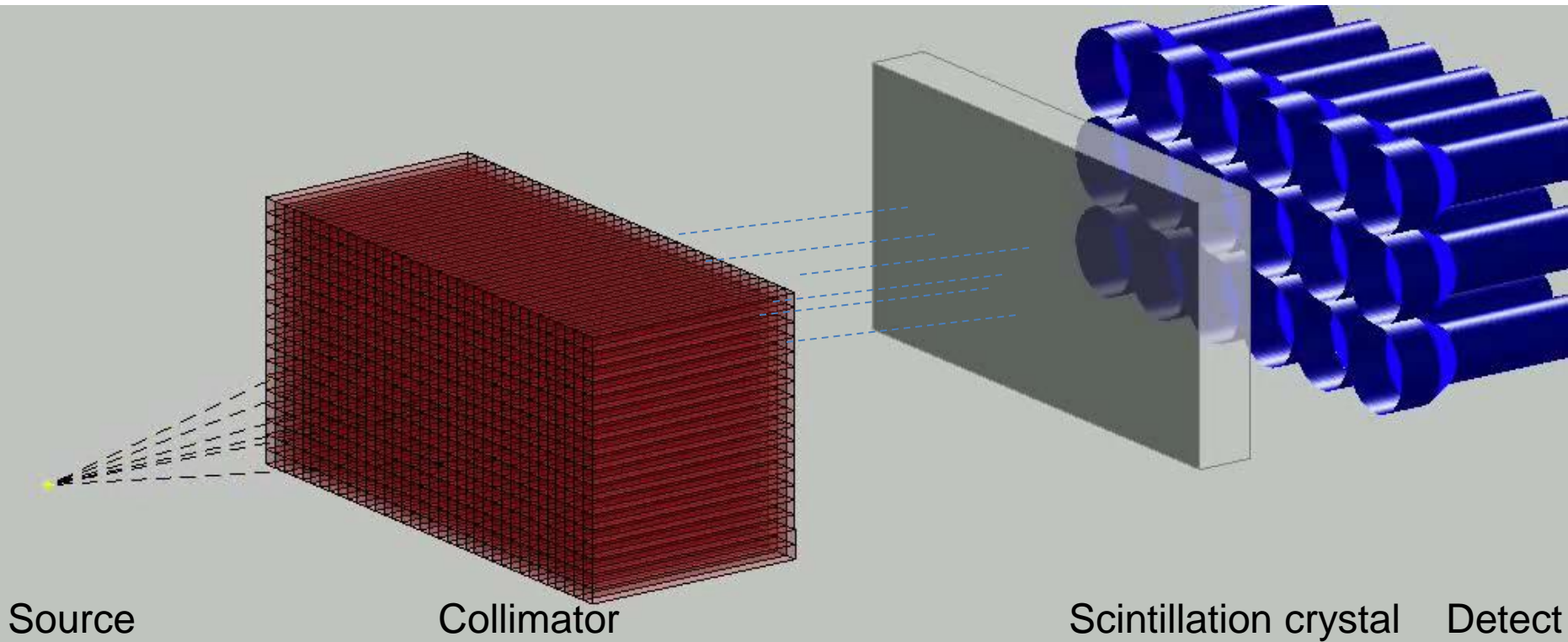




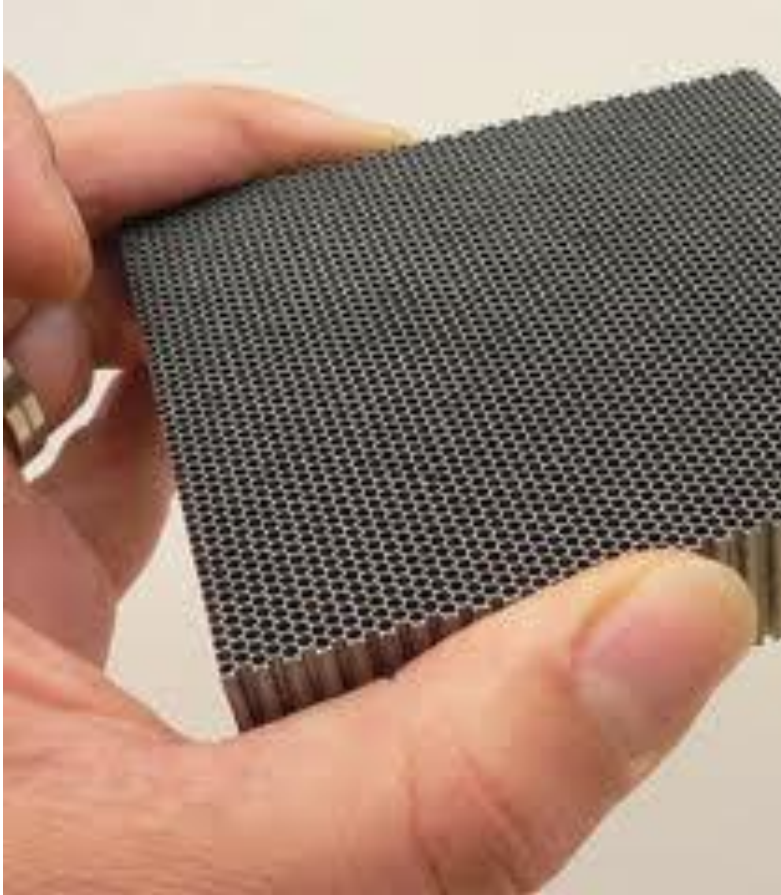
# SPECT Advantages/Disadvantages

- Relatively inexpensive
- Broad experience
- Disadvantages
  - Radiation exposure
  - Preparation of imaging agent
  - Nuclear Regulatory
  - Scanning is slow, low resolution

# SPECT detectors



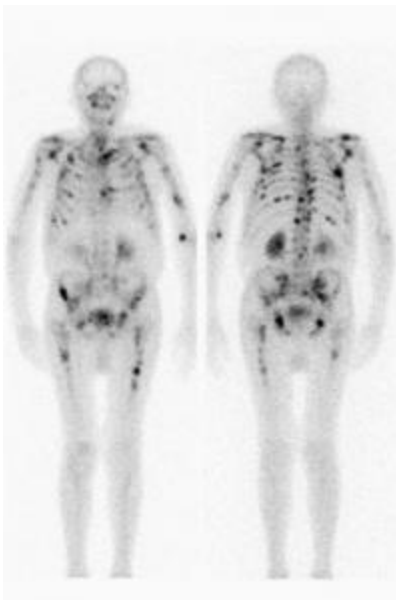
# Collimation cont'd



Collimation reduces the sensitivity and resolution of SPECT by rejecting the majority of events

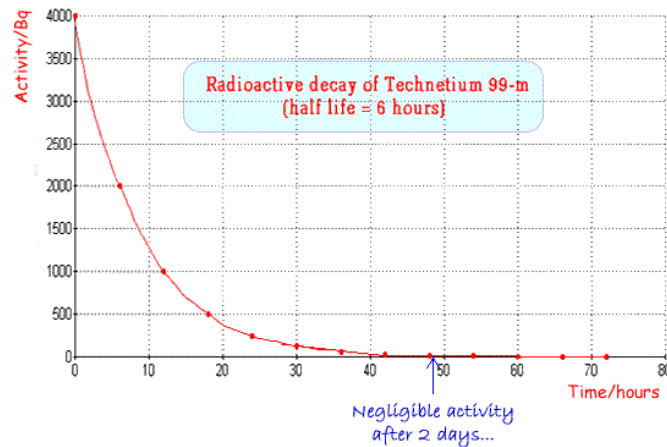
# SPECT Imaging

- Requires conjugation of a radioactive isotope to a compound of interest which is injected into the patient:



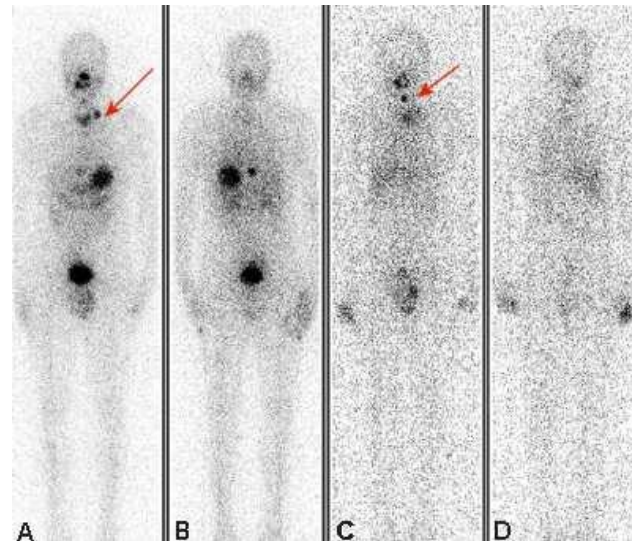
The bone scan:

$^{99m}\text{Tc}$ Technetium-methyl diphosphonate



# SPECT agents for cancer

- $^{99m}\text{Tc}$  MDP Bone Scan
- $^{99m}\text{Tc}$  Pertechnetate (thyroid, salivary gland)
- $^{201}\text{Tl}$  Chloride (parathyroid)
- $^{111}\text{In}$  oxine (WBC labelling)
- $^{131}\text{I}$  Iodine (thyroid)

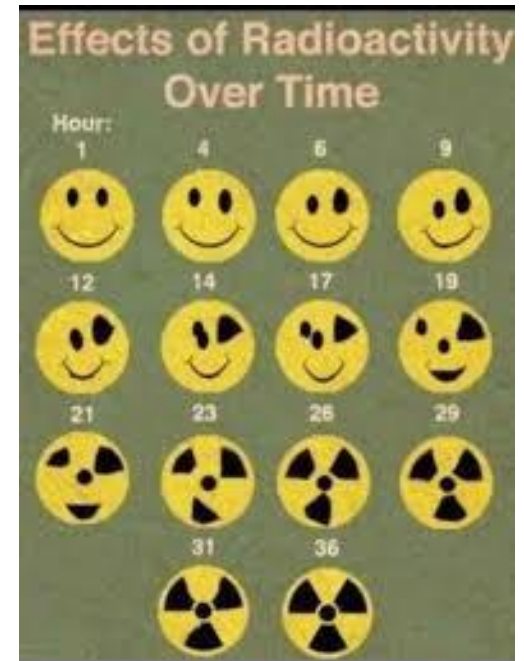
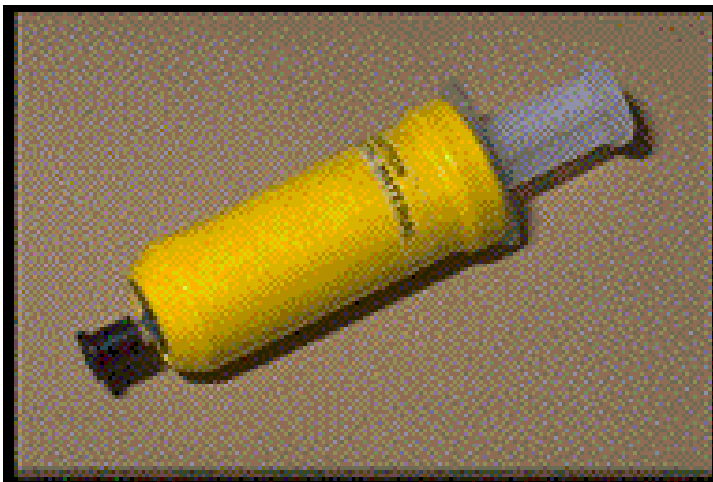


# Hybrid Imaging

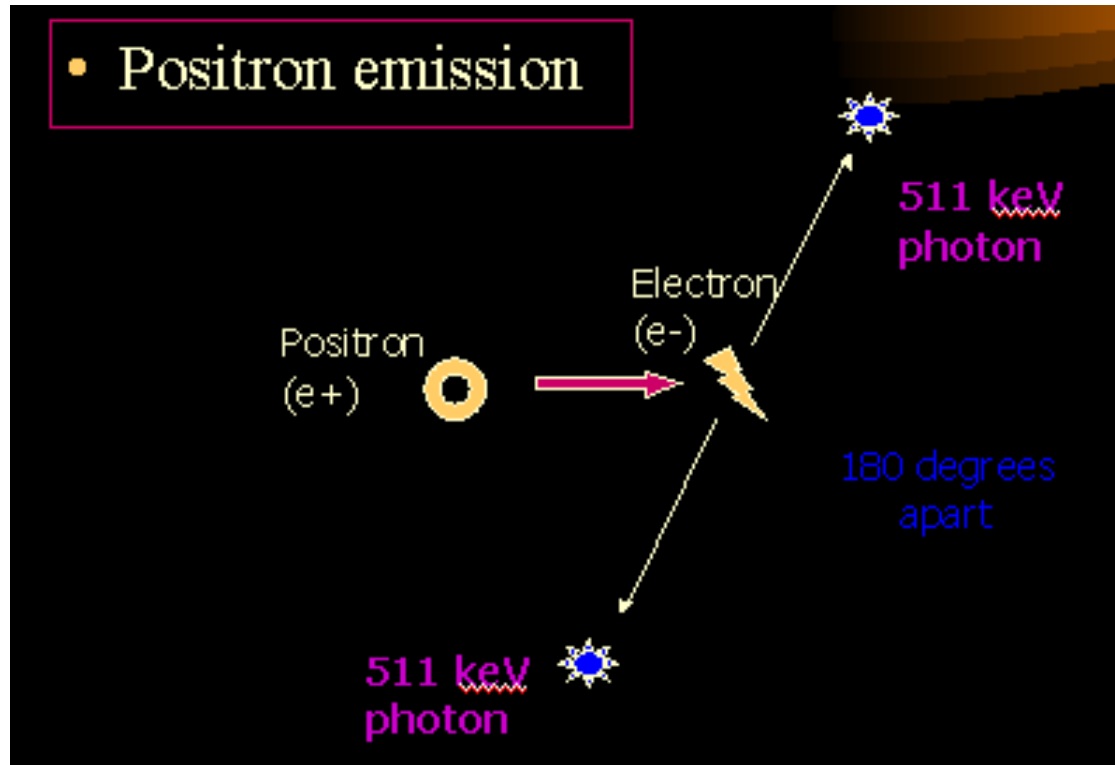


# Safety

- Operator
- Patient



# Positron Emission Tomography

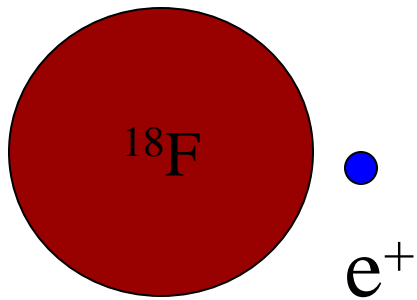


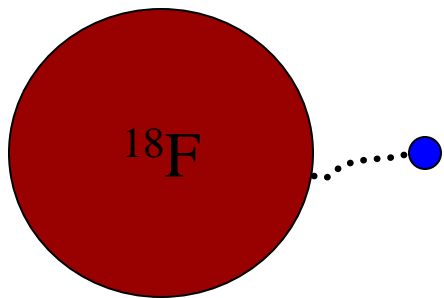


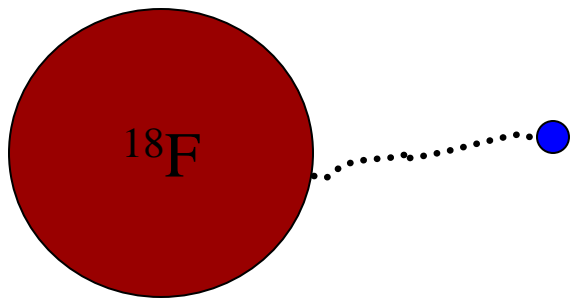
# PET: Advantages and Disadvantages

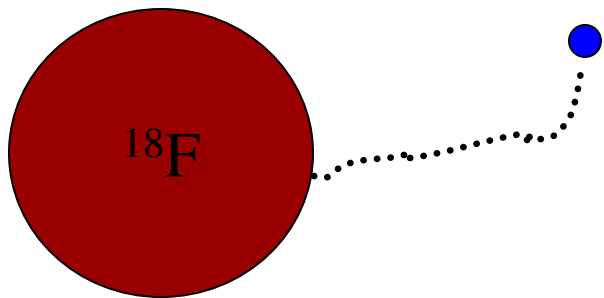
- Highly sensitive
- Metabolic information
- Better spatial resolution than SPECT
- Combined with CT
- ---
- Expense
- Regulatory
- Short half life

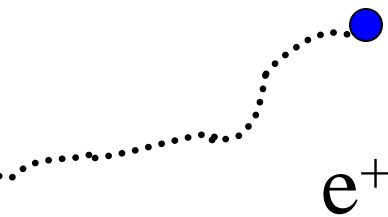
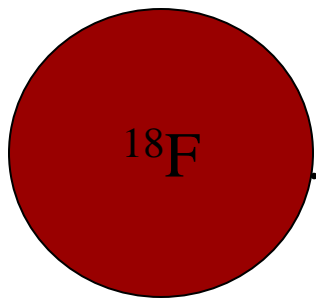
# Positron Emission Tomography

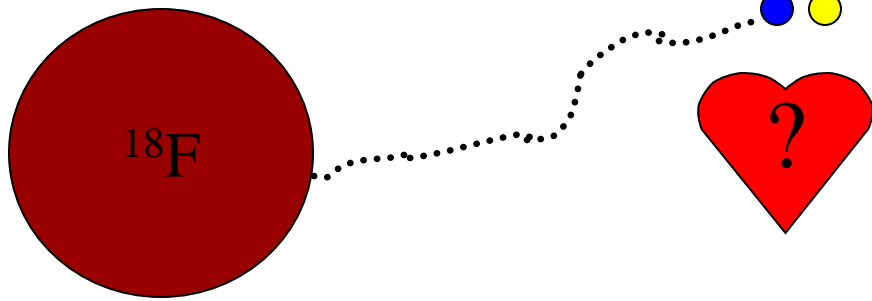


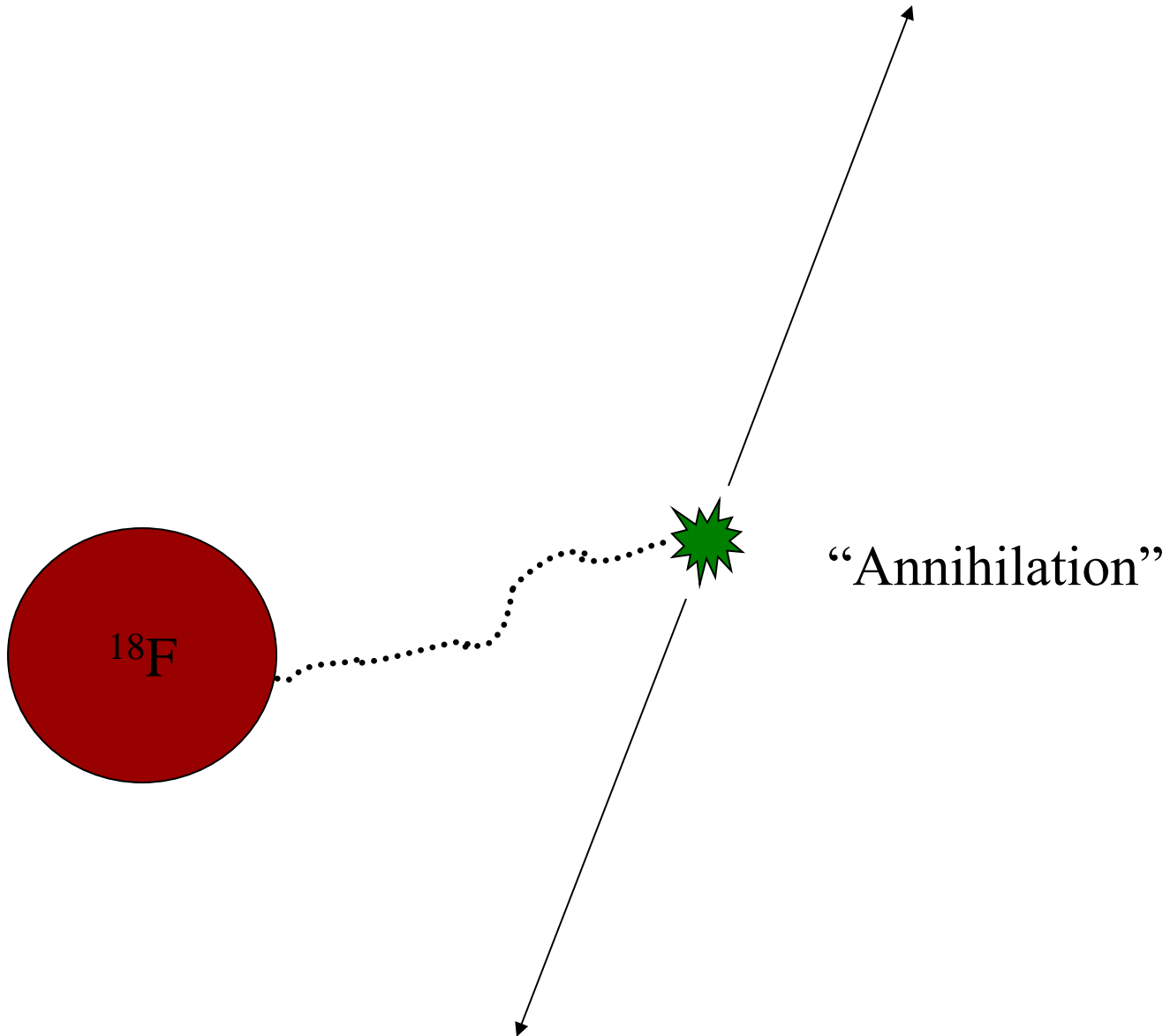




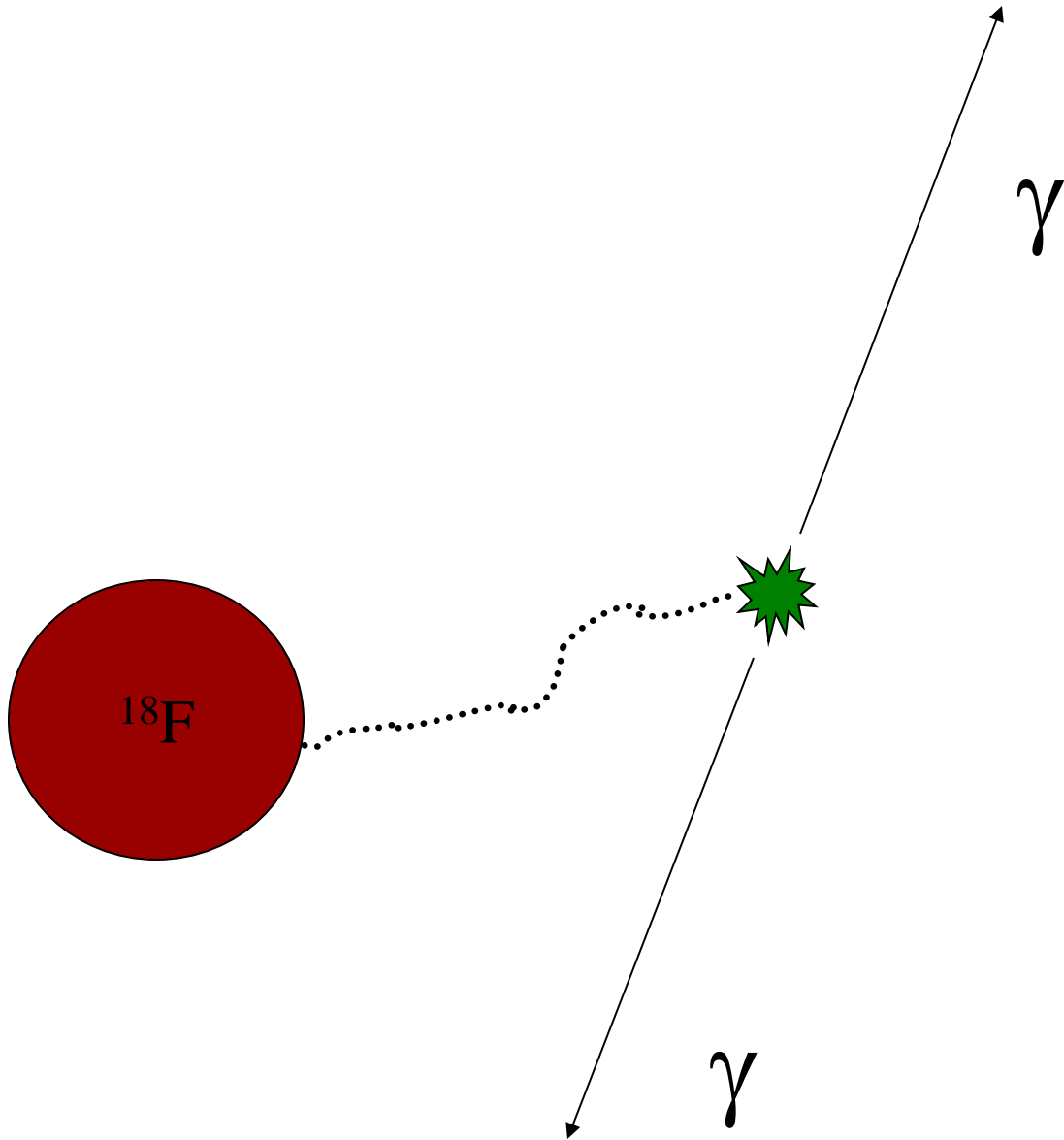




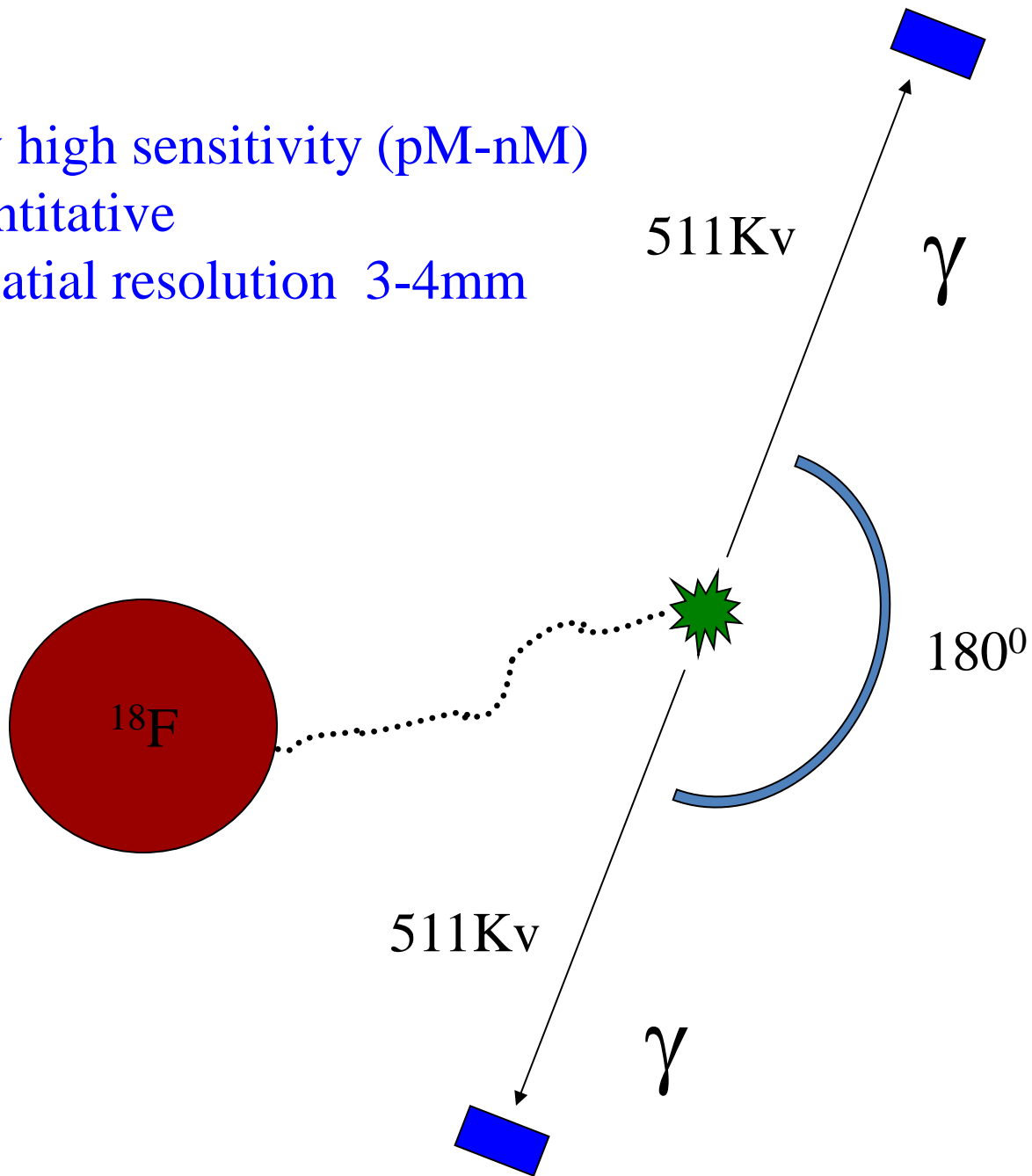








Very high sensitivity (pM-nM)  
Quantitative  
± Spatial resolution 3-4mm



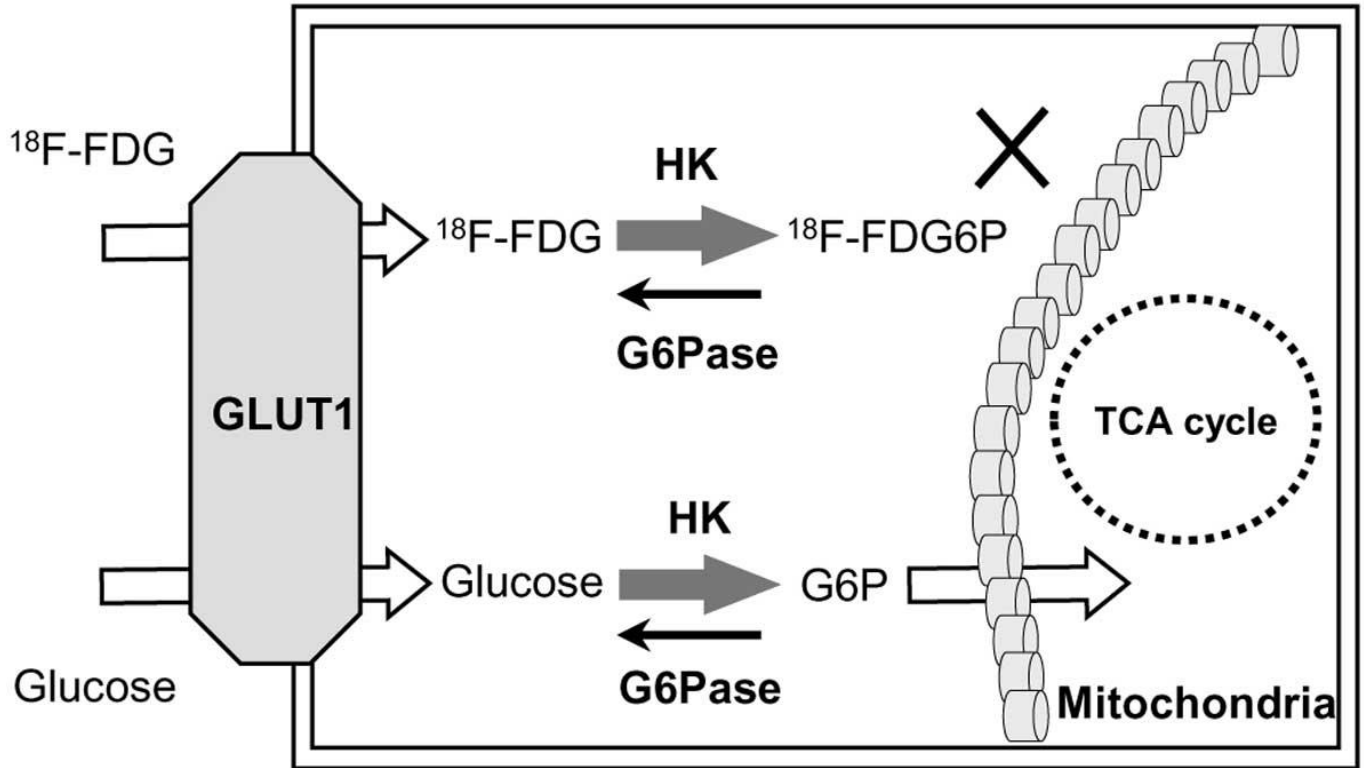
# F-18 Deoxyglucose



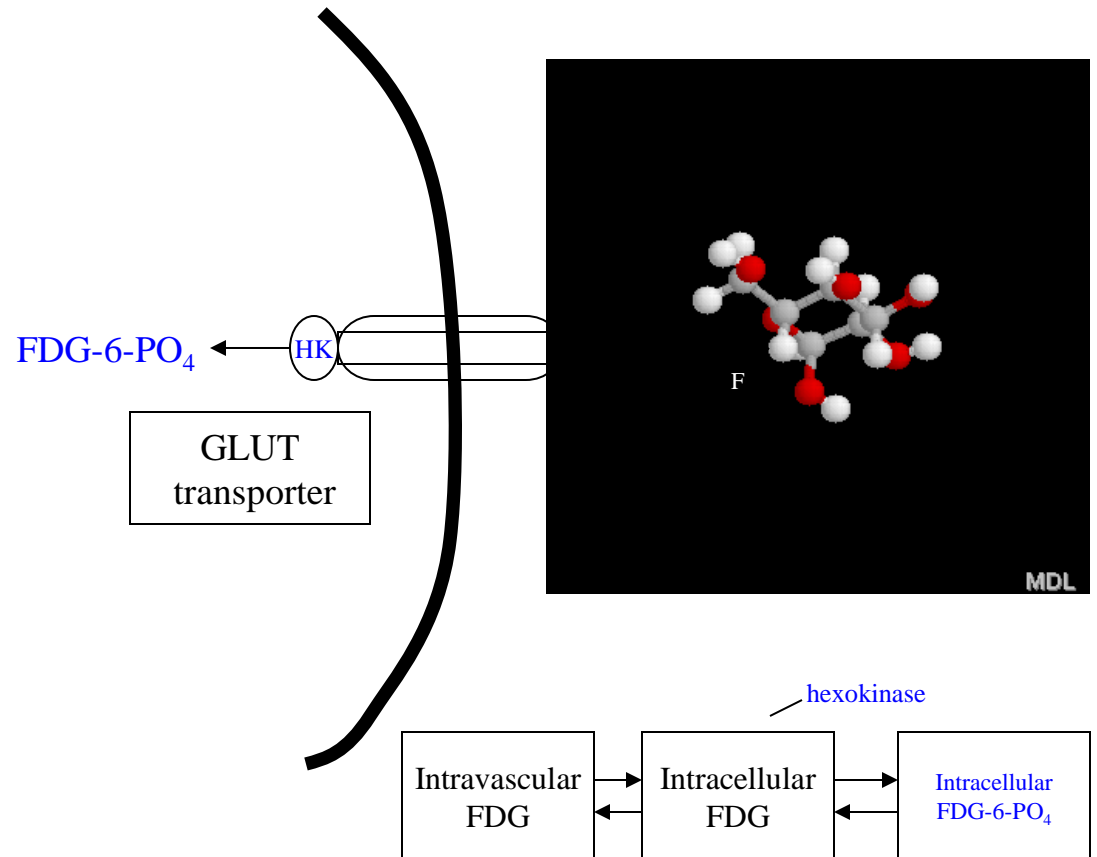
Otto Warburg



Lou Sokoloff

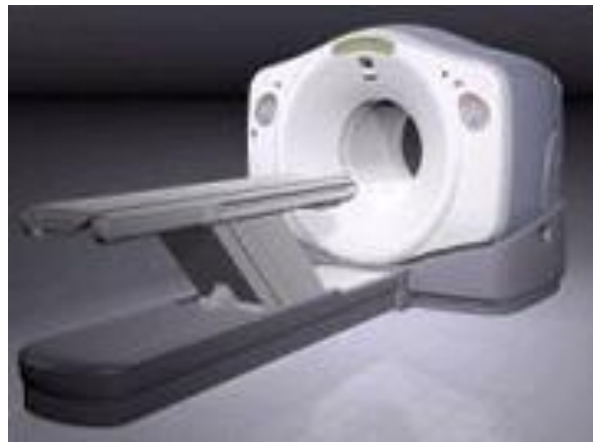


# $^{18}\text{F}$ FDG PET Imaging



# Facilitate Advanced Imaging Technology

- Positron Emission Tomography
  - PET-CT Device



PET Image

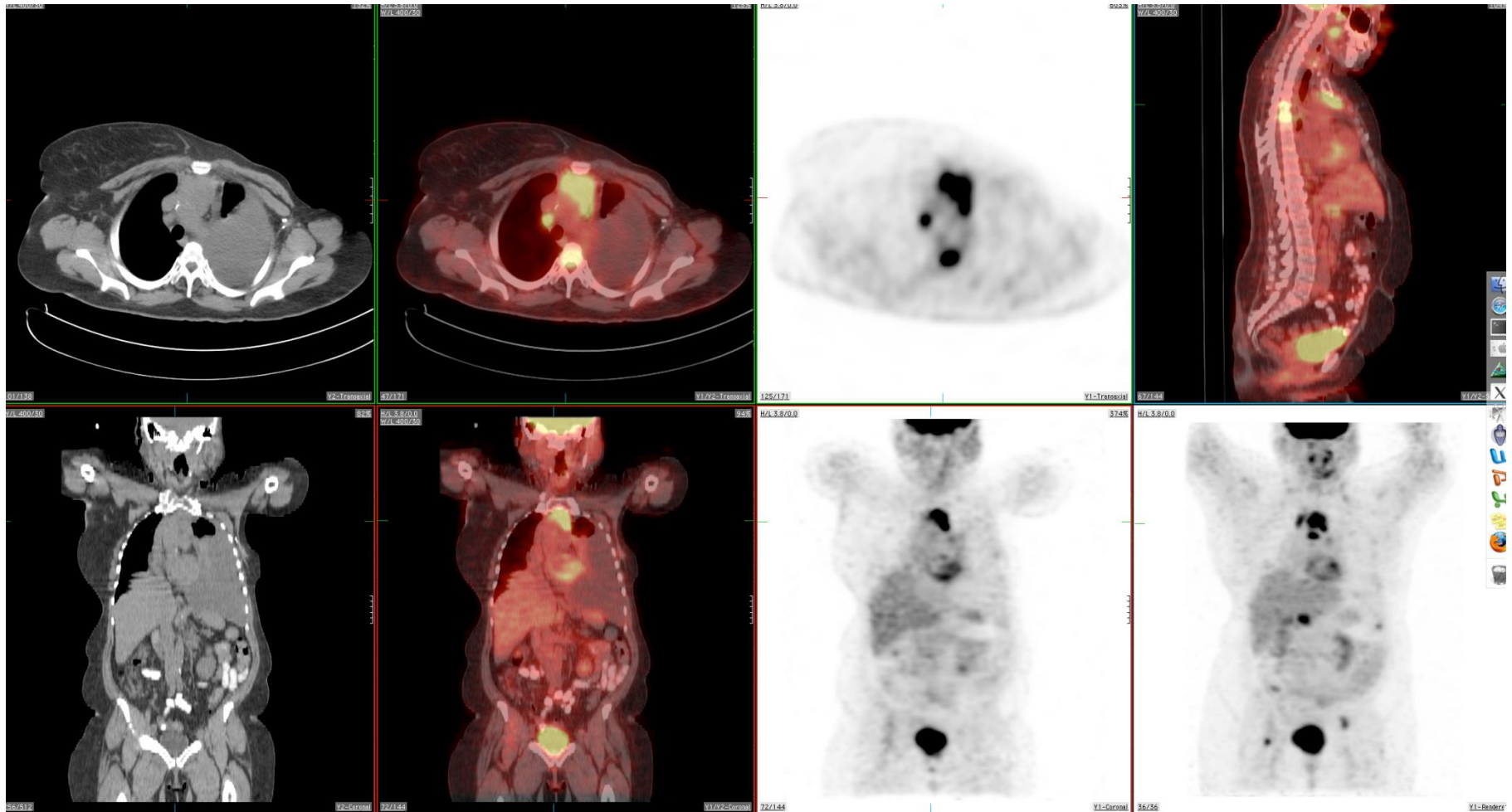


CT Image



Discovery™ LS

# Mediastinal and spine metastases (breast)



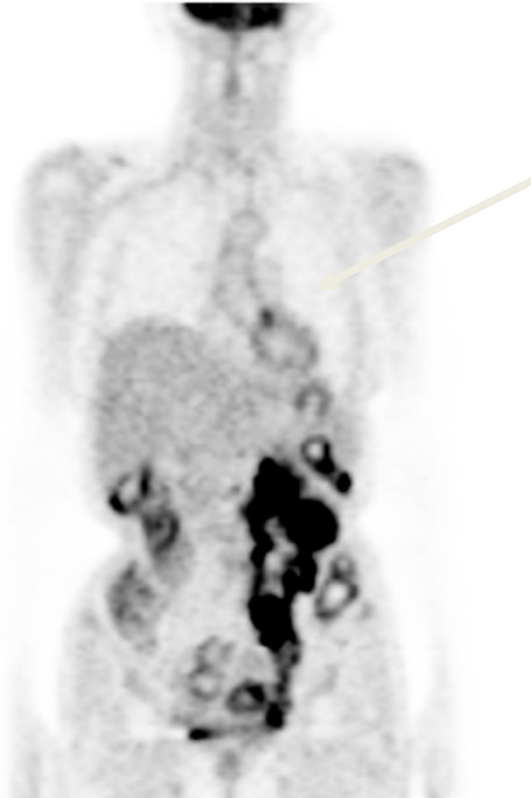
Metastatic Breast Cancer

# Notable PET Agents

- Sodium Fluoride: Bone target
- Fluorothymidine: Cellular Proliferation
- Fluoroestadiol: Estrogen receptor
- Fluorocholeline: Membrane Turnover
- Fluoromiso: Hypoxia
- Florbetaben: Amyloid (Alzheimers)
- Zirconium Herceptin: labeled antibody
- Zirconium Oxine: Cell labeling



**Non-attenuation  
corrected**



**Attenuation  
corrected**



# PET Imaging

- Positron emission tomography (PET) has the advantages of :
  - High energy photon imaging
  - High Sensitivity, Moderate Specificity
  - The ability to correct for attenuation
  - No need for collimation
  - Resolution is still limited

# Summary of Cancer Imaging

Presentation	Resolution	Sensitivity	Cost (low-hi)
CT	CT	PET	US
MRI	MRI	SPECT	CT
US	US	US (microbubble)	SPECT
SPECT	PET	MRI	MRI
PET	SPECT	CT	PET

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# General Guidelines

- Overall “workhorse” for oncology: CT
- Specialty cancers: brain, liver, prostate: MRI
- Problem solving (e.g cyst vs. solid): US
- Bone mets: SPECT
- Metabolic activity: PET



# Imaging of Cancer:

<http://mip.nci.nih.gov>

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