

Imaging of cancer



Imaging of Cancer:

How various imaging modalities work

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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

Scanners

They all look the same!



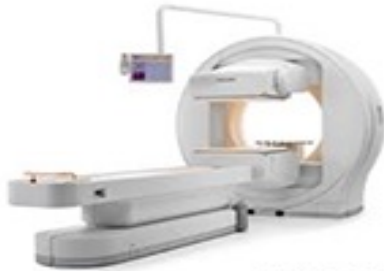
CT



MRI



US

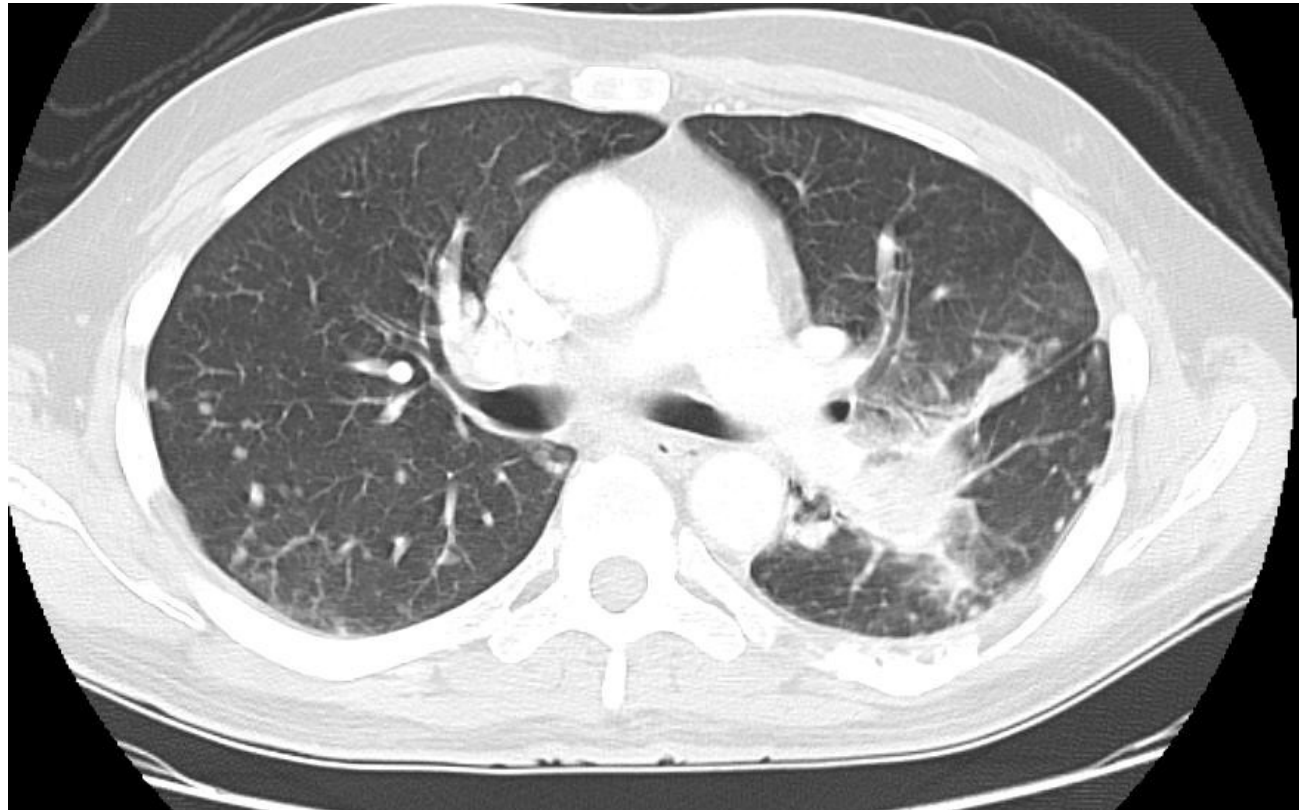


SPECT

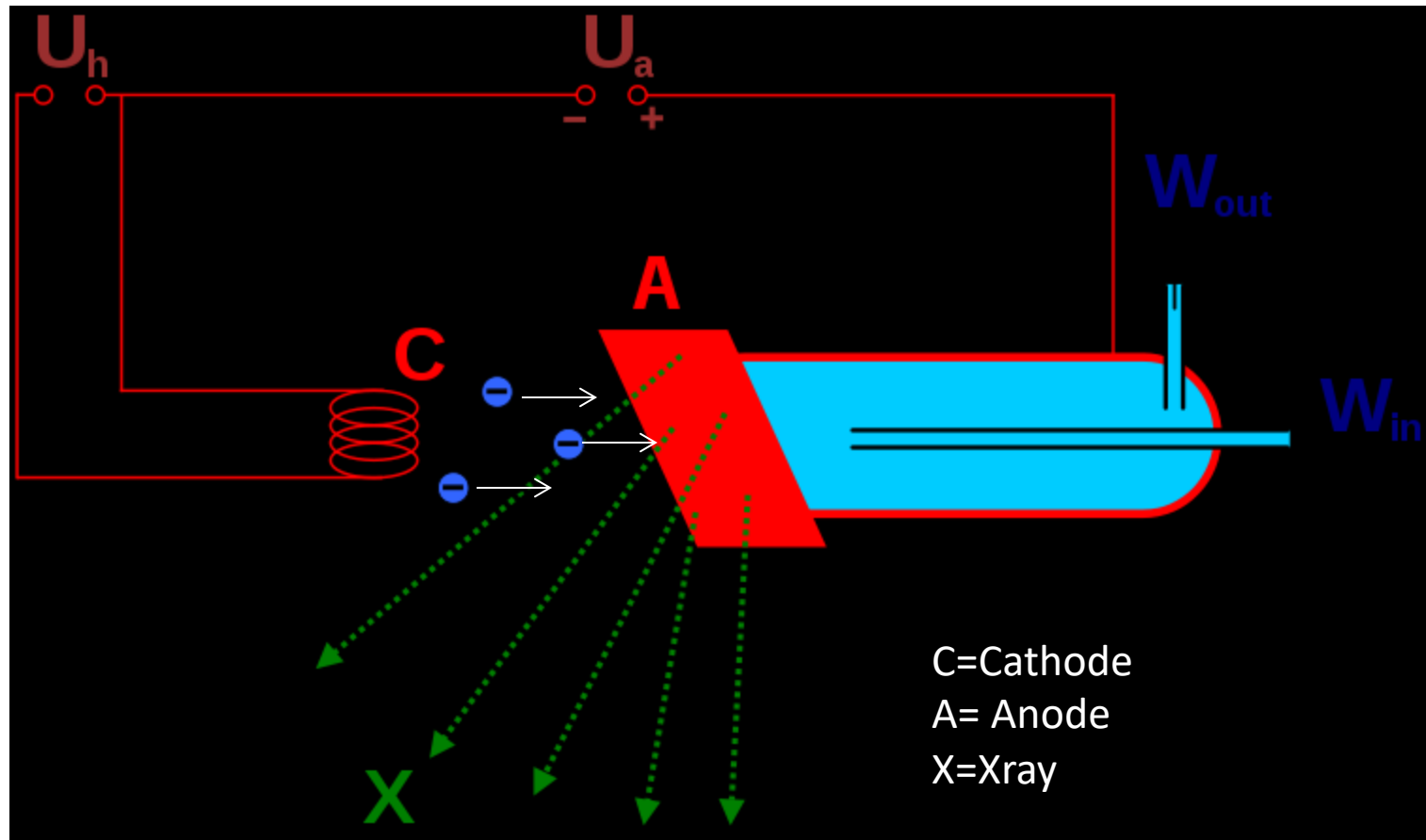


PET

Computed Tomography

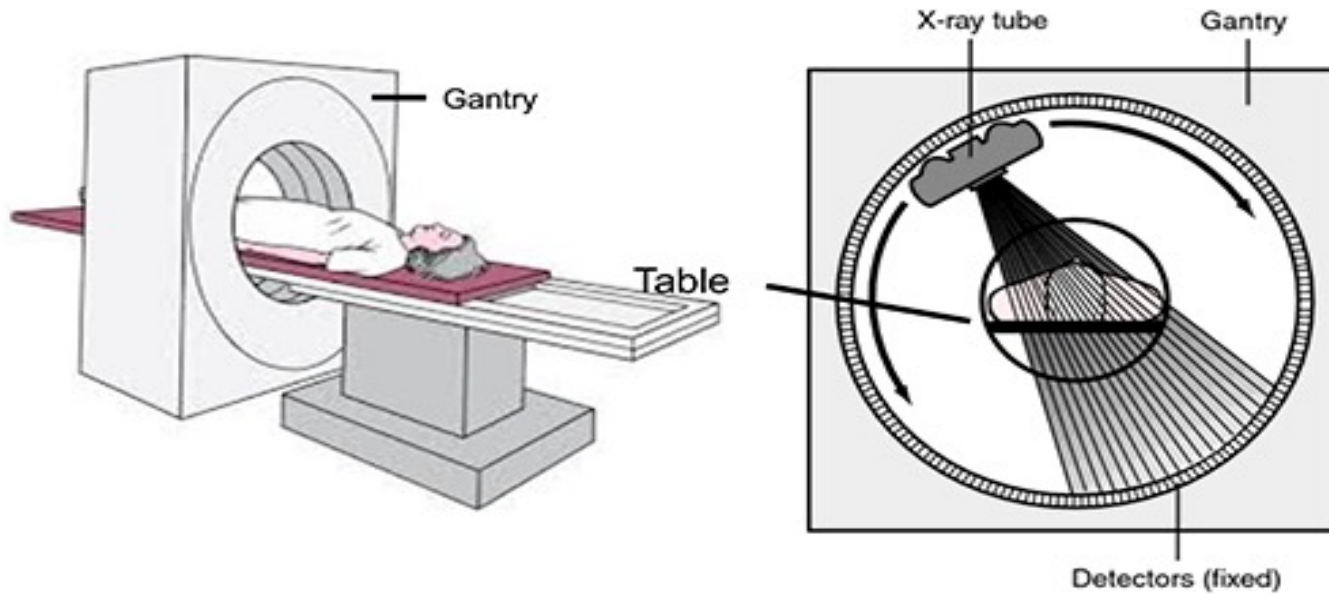


X-ray production: cathode ray tube



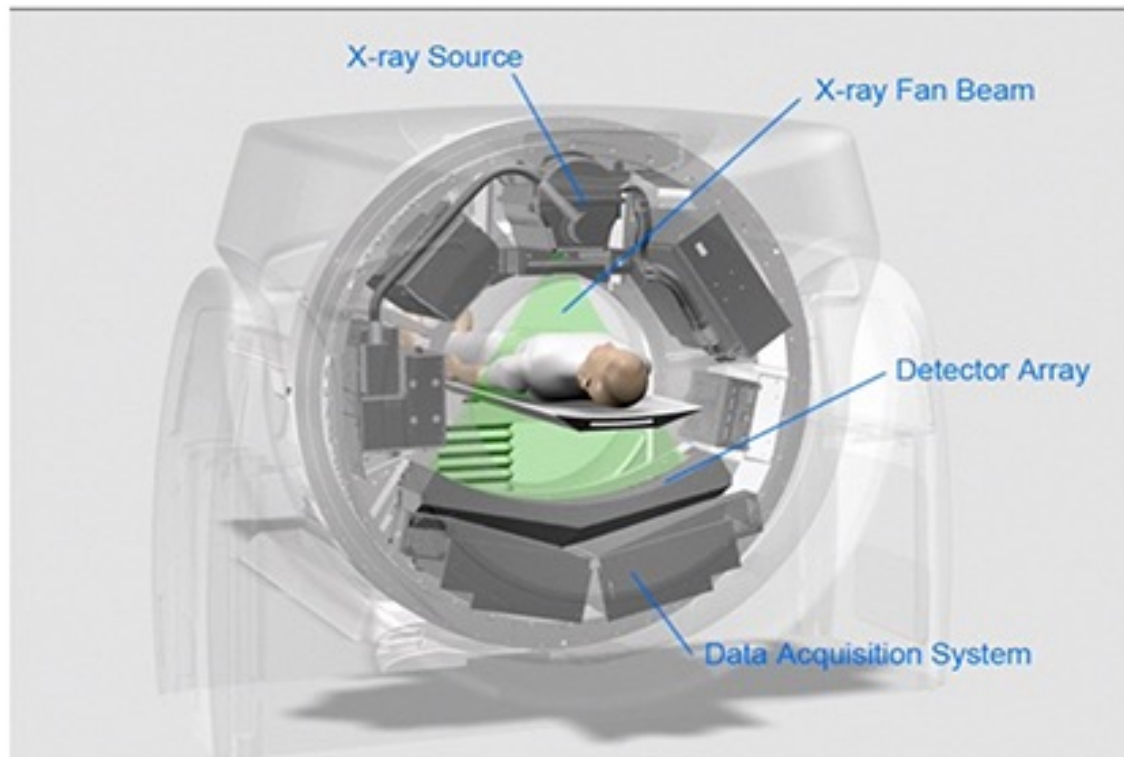
Basics of CT

Basics of CT



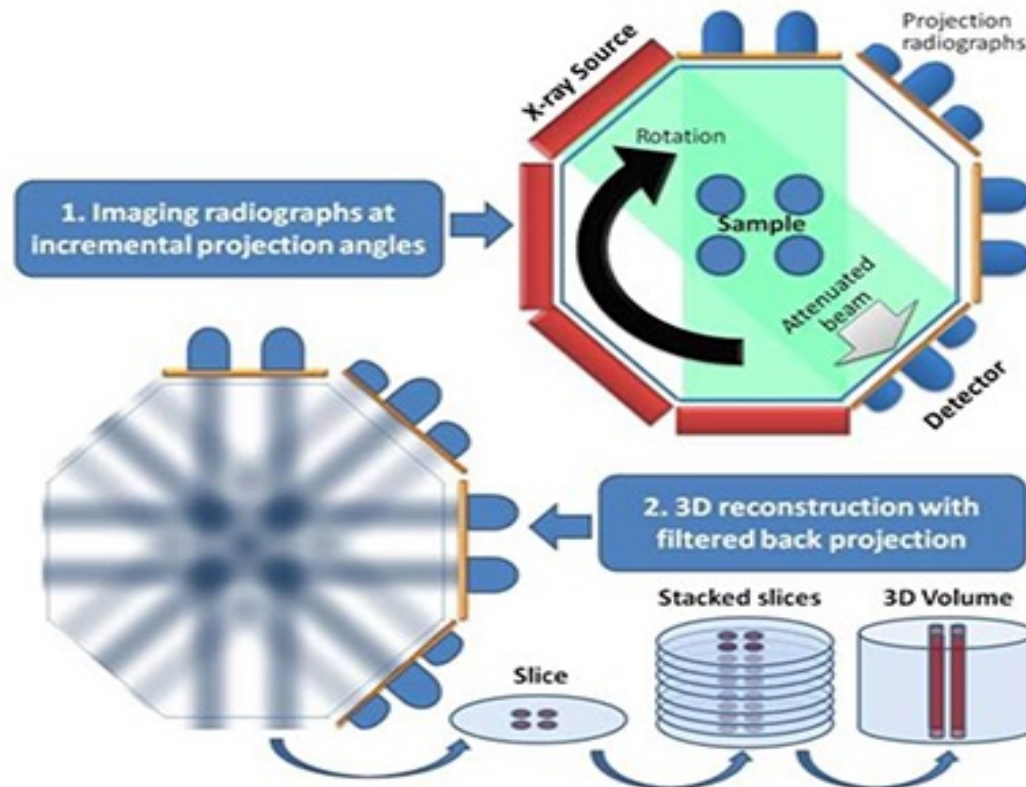
CT scanner

Cross section of a CT Scanner



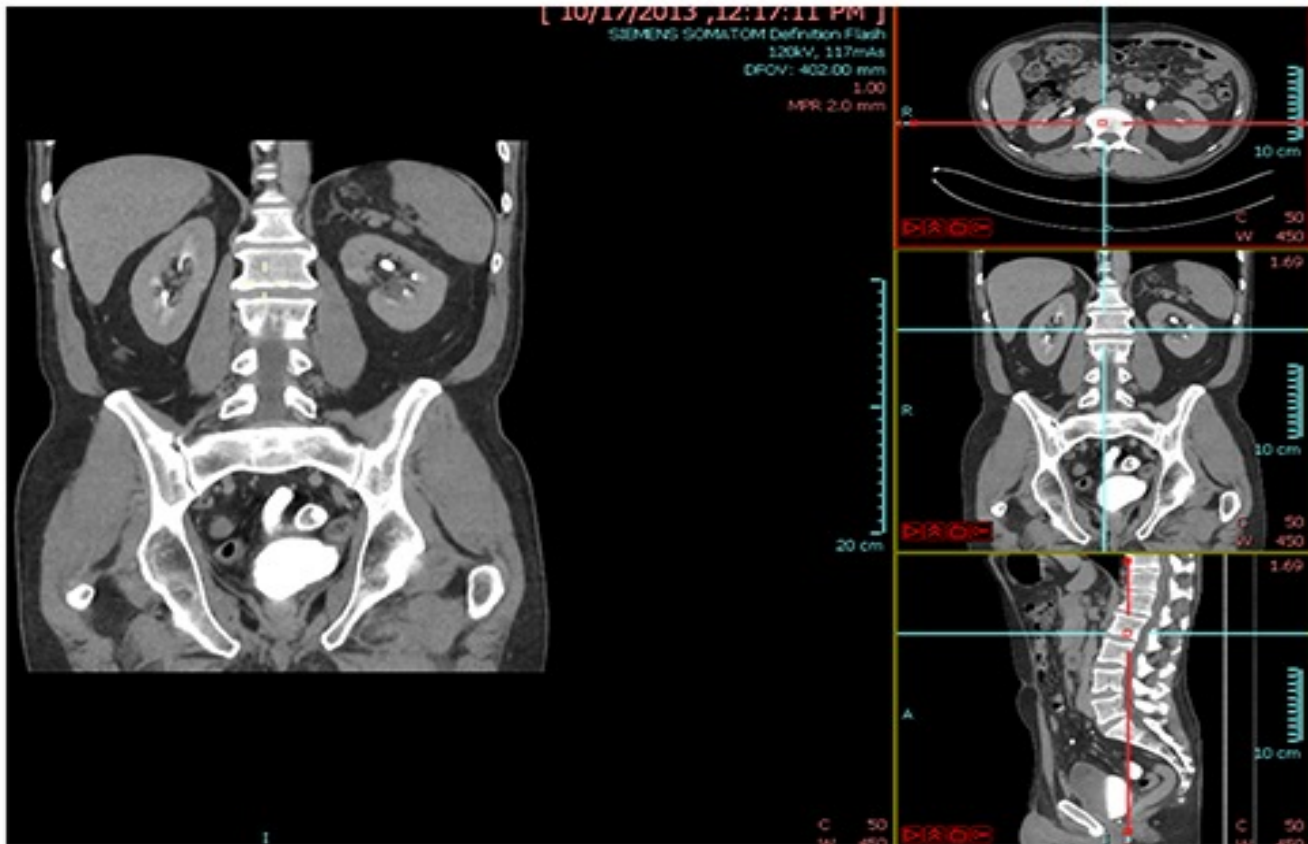
Filtered back projection

Filtered Back Projection



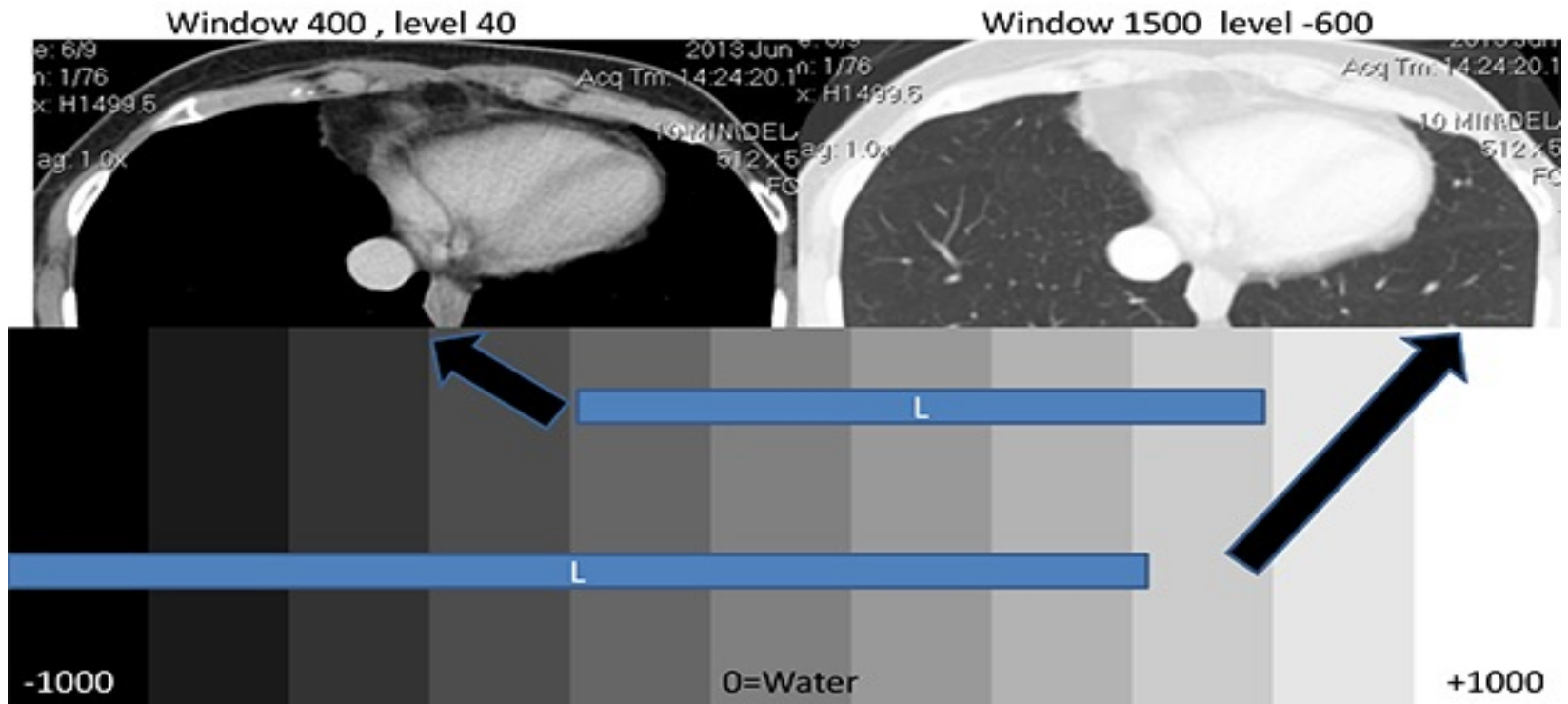
Volume CT imaging

“Volume” CT imaging



Windowing a CT

“Windowing” a CT: The value of a digital image



Advantages of CT

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

Disadvantages

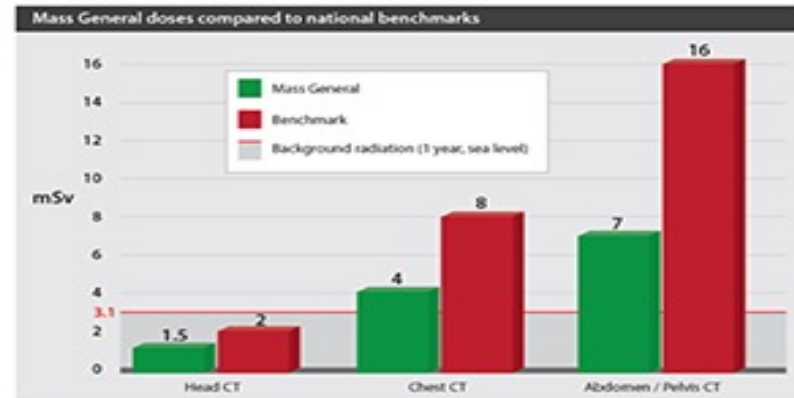
- **Radiation**
- Often requires iv contrast media
 - Allergic reactions (minimal)
 - Kidney damage (only in high risk patients)
- Anatomic information only

Radiation reduction

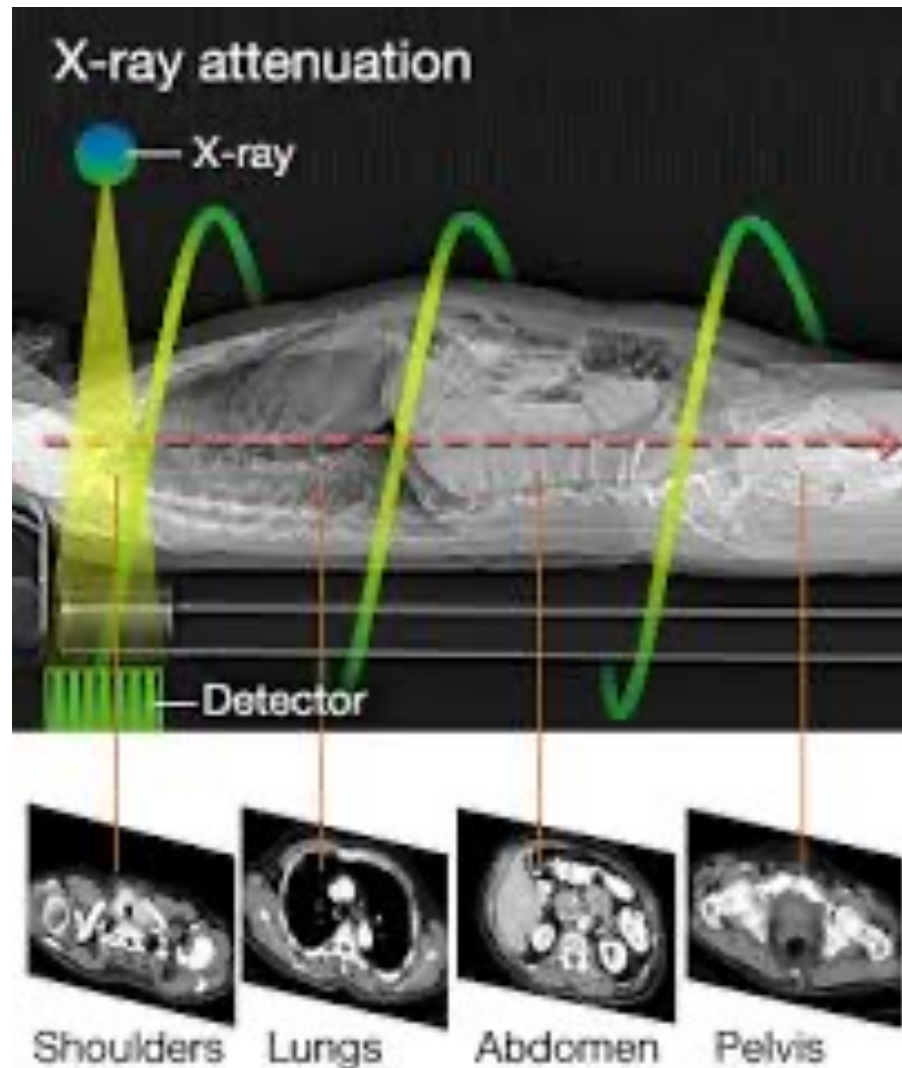
Radiation Reduction on CT



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

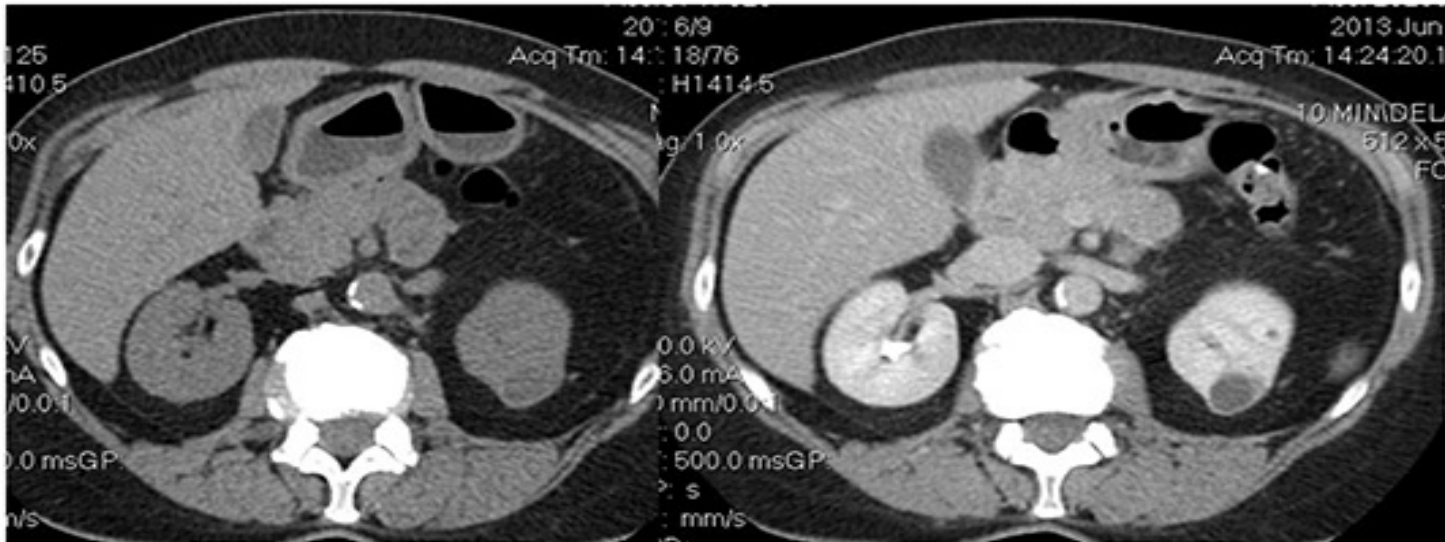


Attenuation differences thru the body mean less radiation for some regions



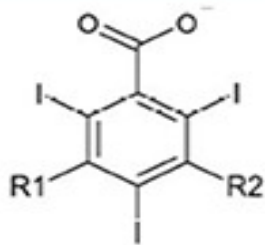
Contrast media

Iodinated Contrast Media

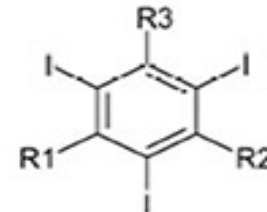


Structures

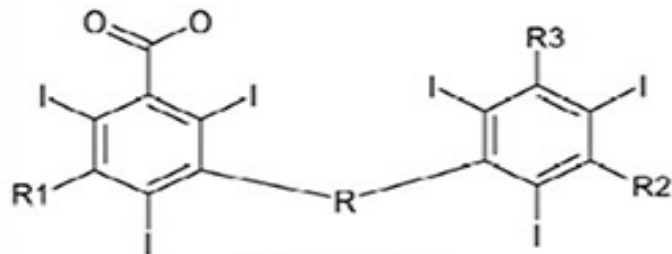
Basic Structures of Contrast Media



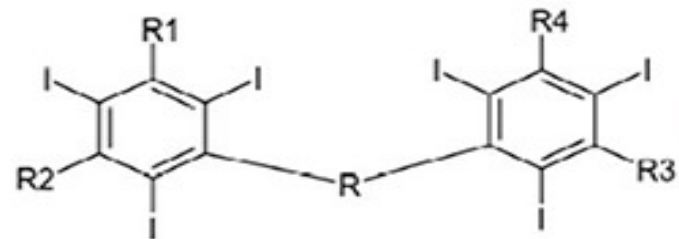
Ionic Monomer



Nonionic Monomer

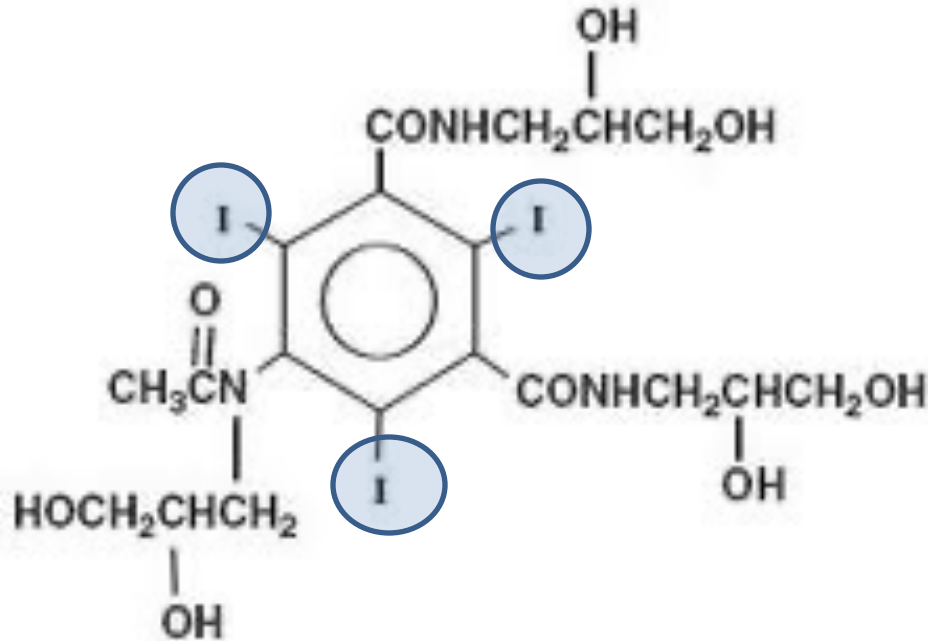


Ionic Dimer



Nonionic Dimer

Non ionic Iodinated Contrast



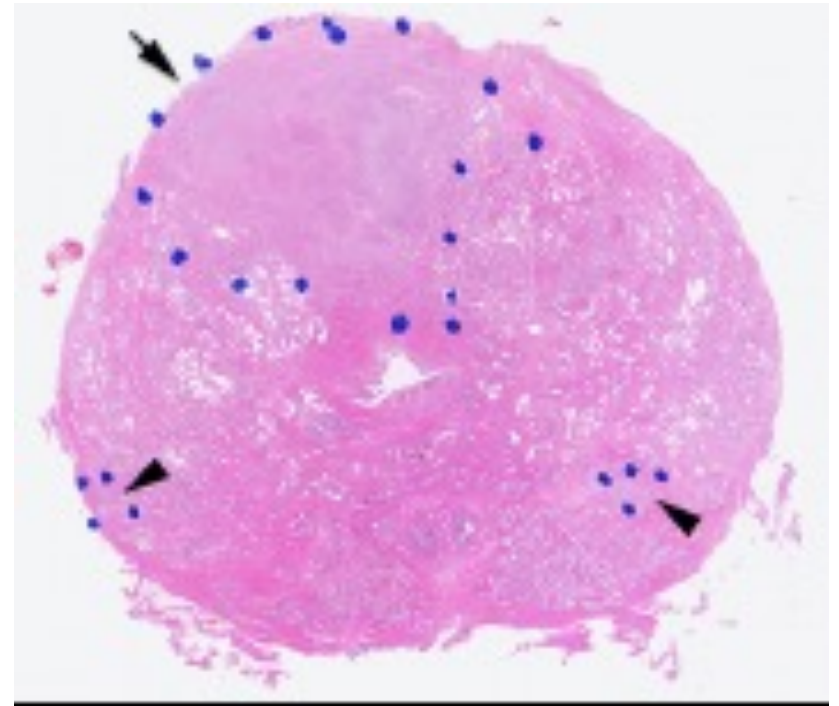
Typical dose 30-45 Grams of Iodine!!!

Iodinated contrast

Iodinated Contrast

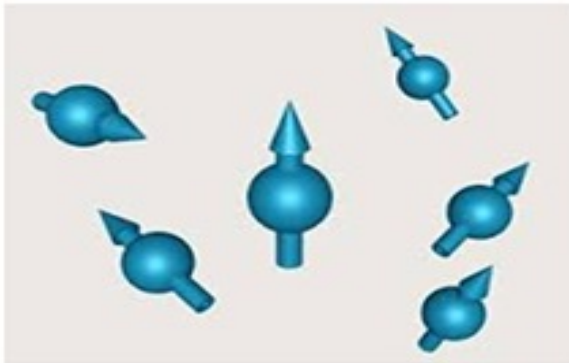


Magnetic Resonance Imaging

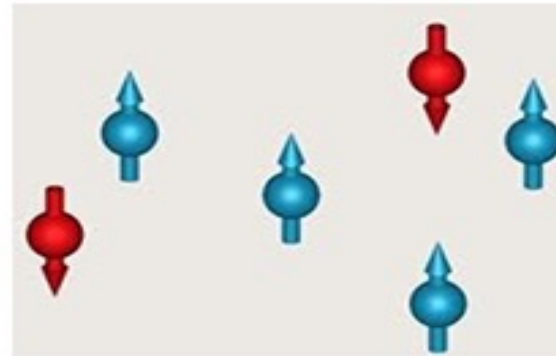


MRI physics

MRI Physics 101

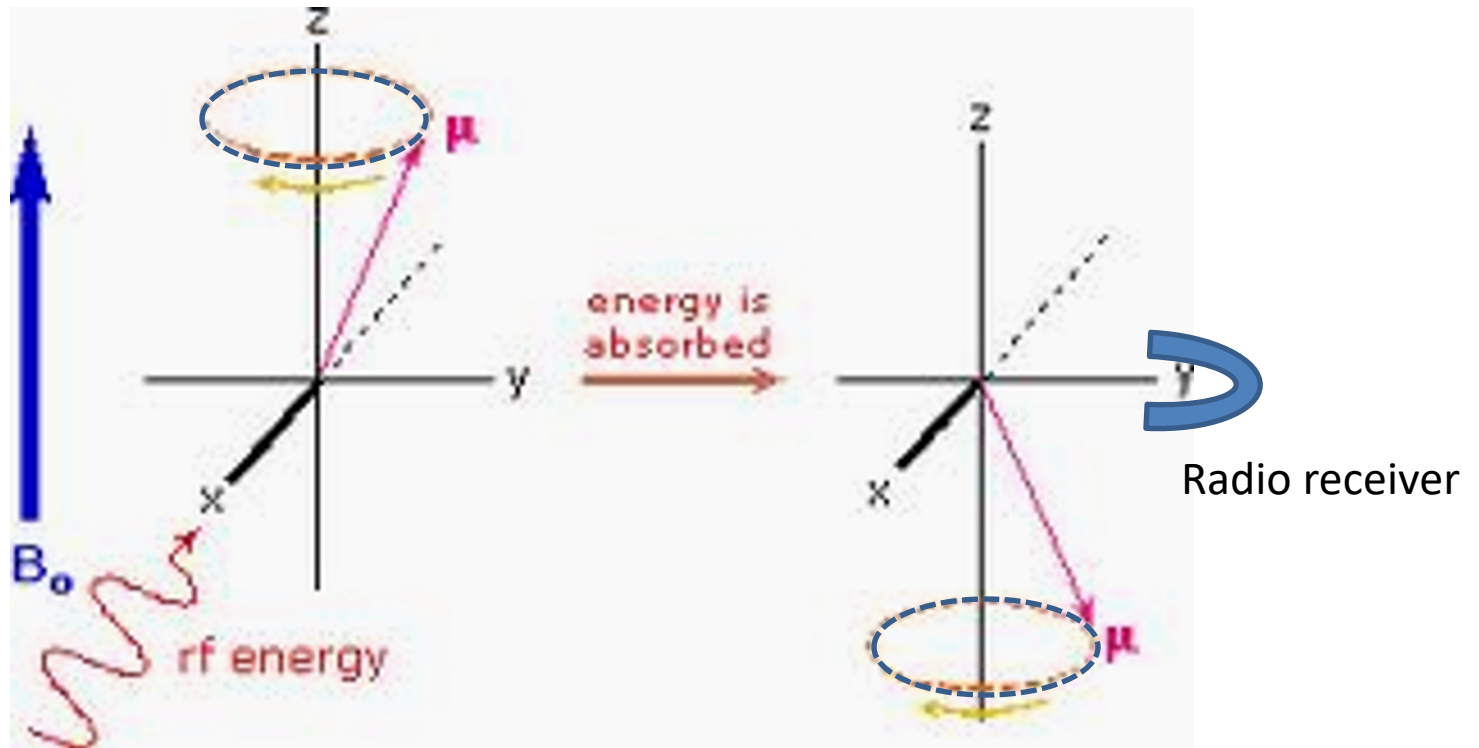


Protons in space: no field

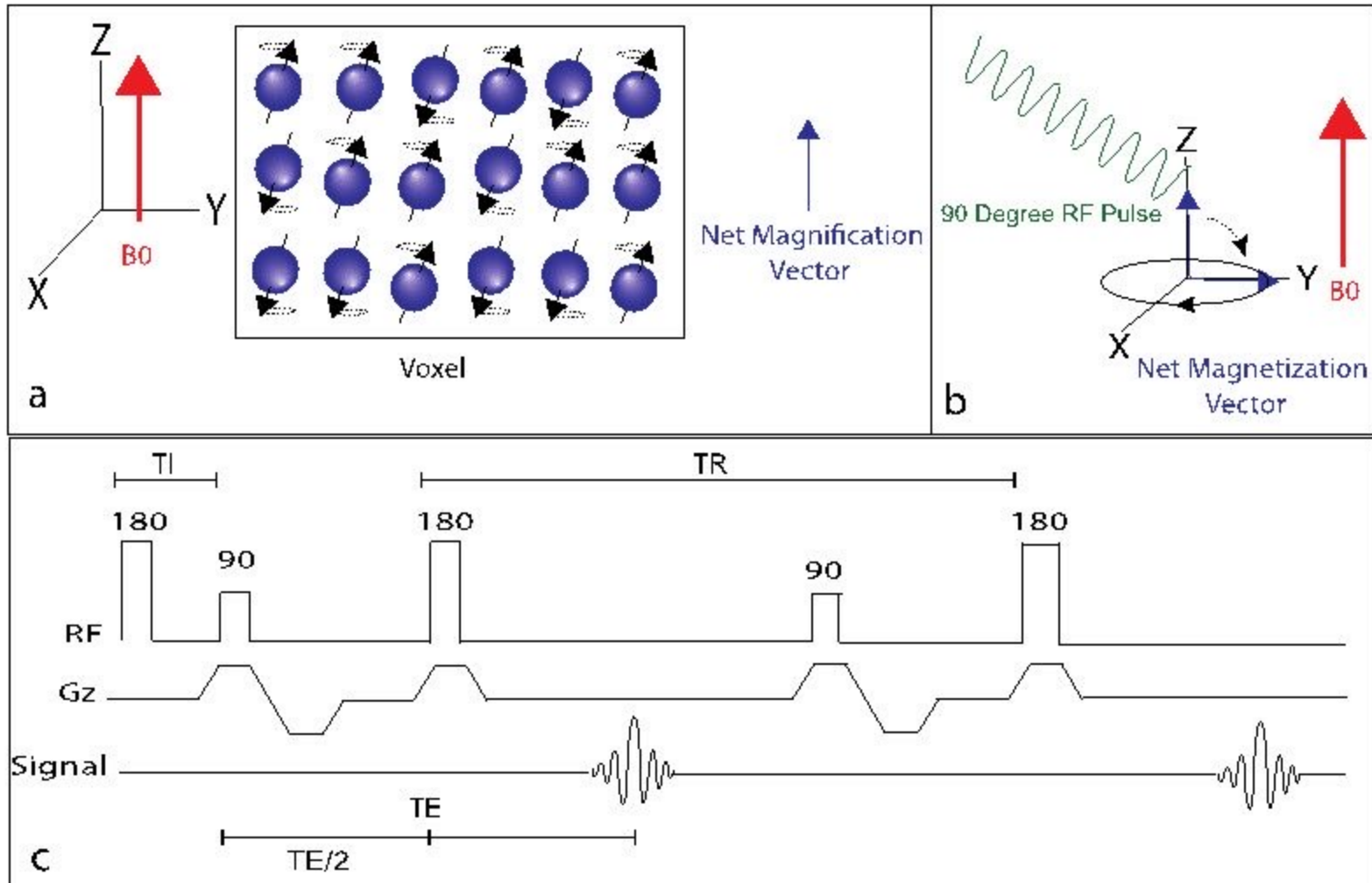


Protons in magnetic field

MR Physics

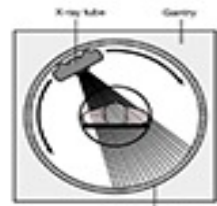


Summary

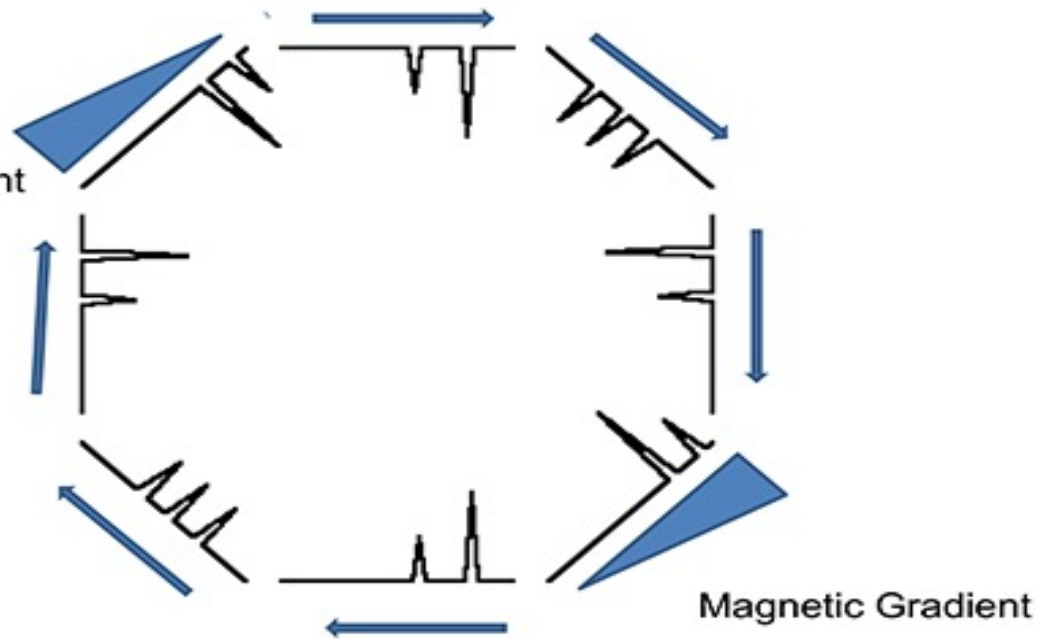


MRI image

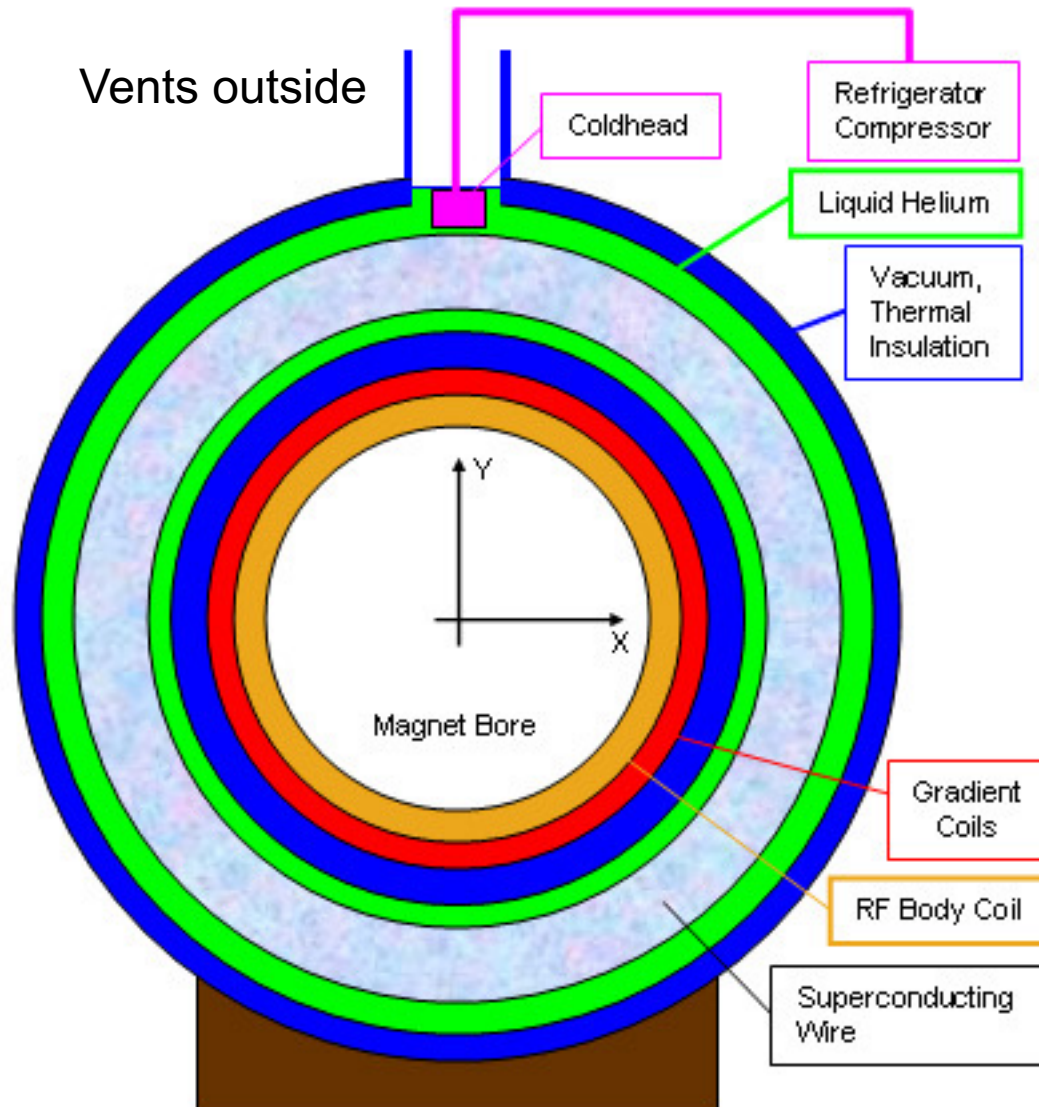
Creating an MR Image:
No detectors! Just antennas (coils)



Magnetic Gradient



Anatomy of an MRI



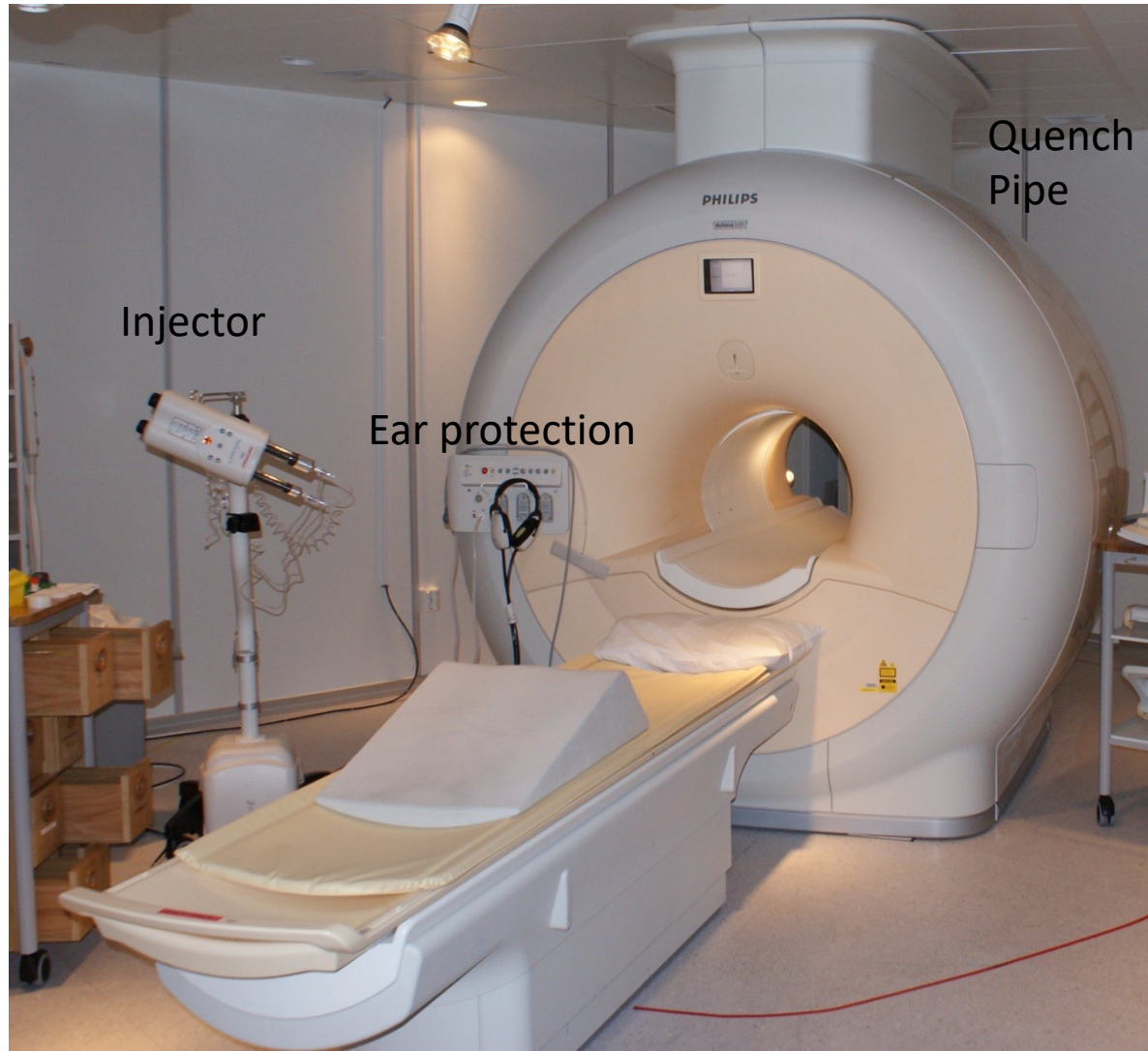
MRI Advantages

- 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

Safety issues in MRI



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



MRI

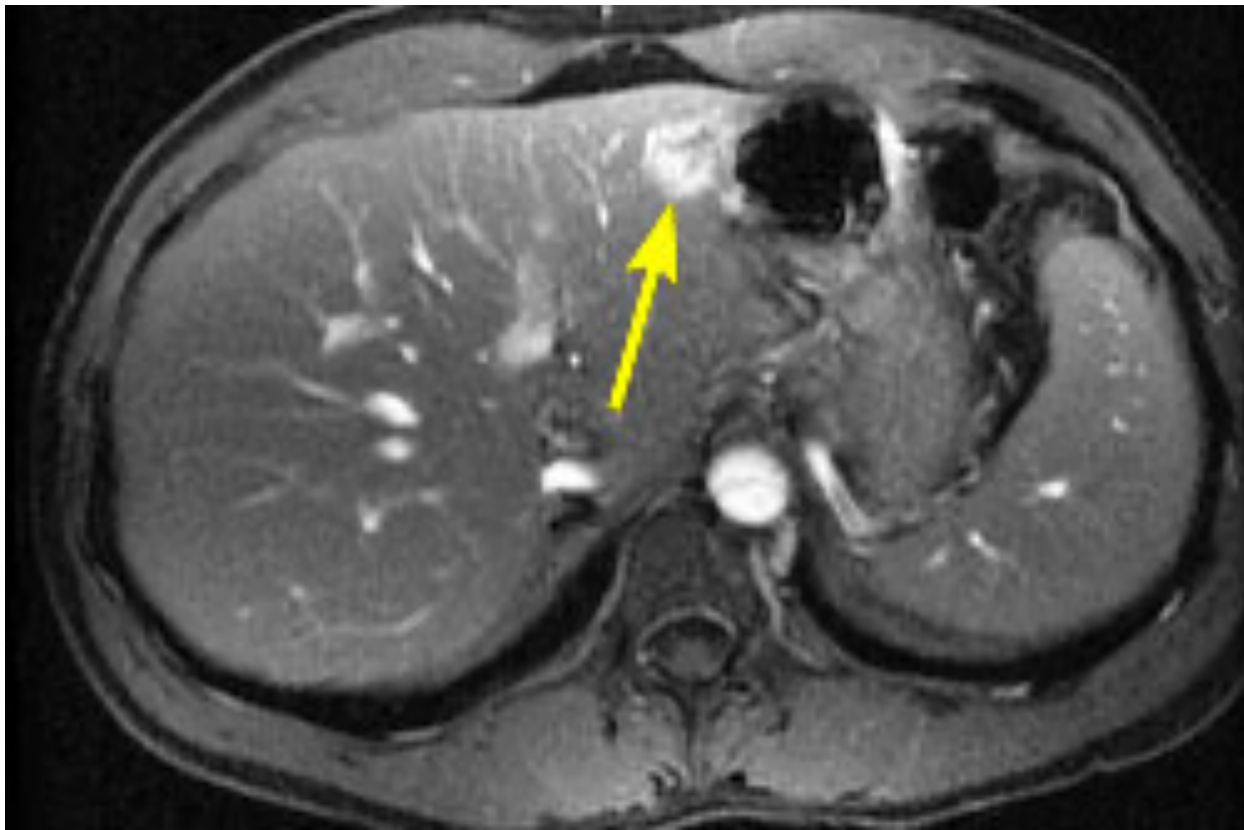
O2 Tank, "Missile"



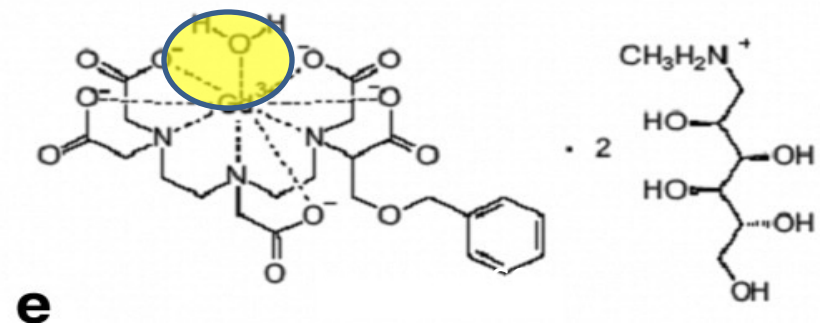
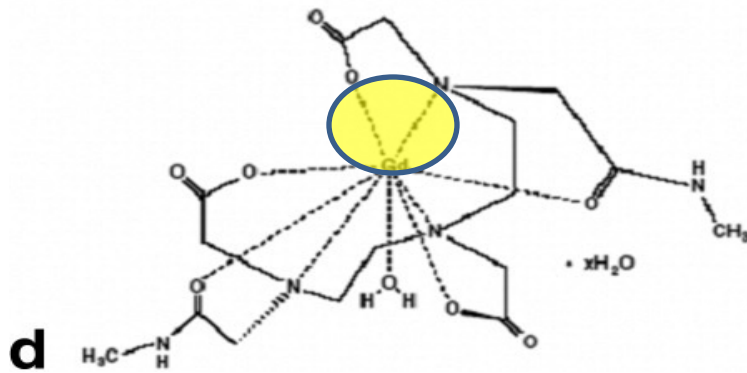
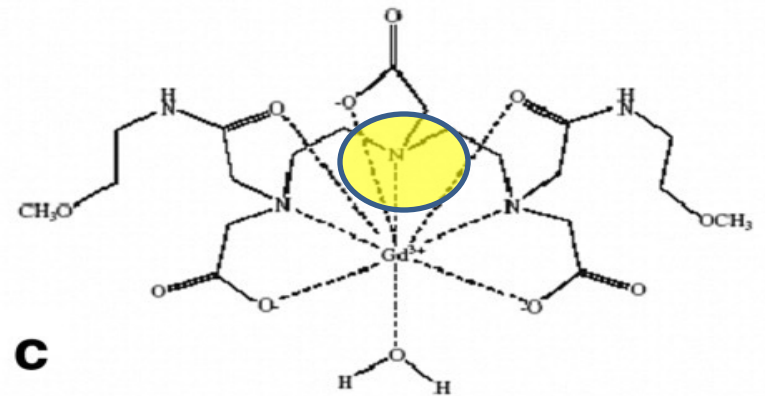
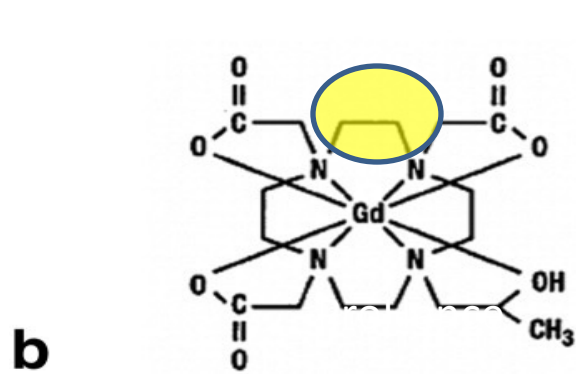
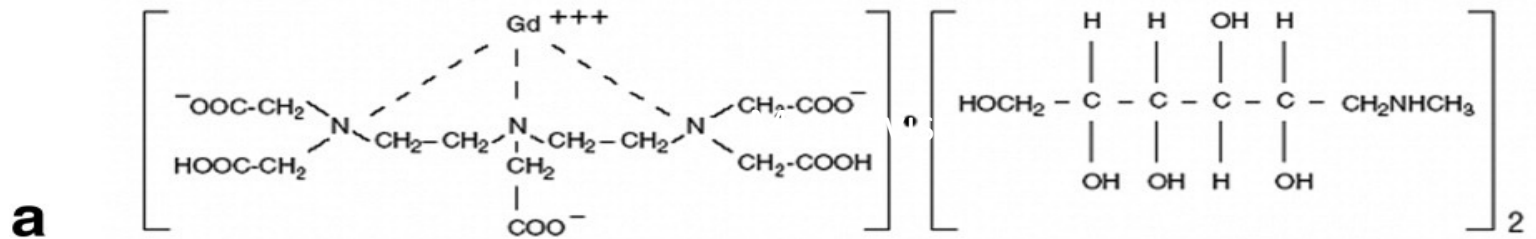
An Oxygen tank can become an Airborne torpedo in an MRI



Value of Contrast Media



Contrast agents



Contrast table

Extracellular Gd-CM	Type	Thermodynamic stability constant	Conditional Stability	Amount of excess chelate (mg ml ⁻¹)	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

Fibrosis

Nephrogenic systemic Fibrosis



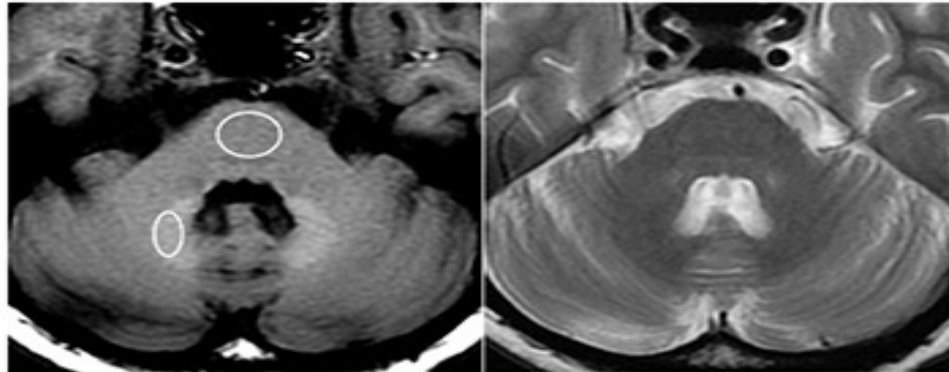
Patient with very poor renal function received multiple linear Gd injections for MRI.

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.

Residual gadolinium

Residual Gadolinium!



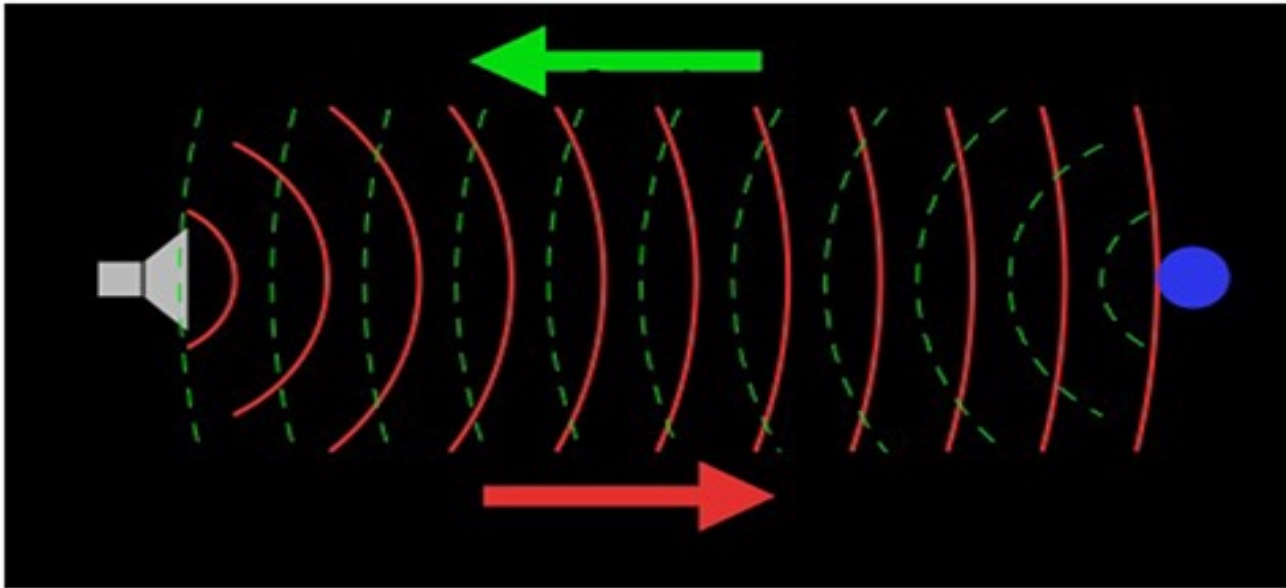
Extravascular Gd:CM	Type	Thermodynamic stability constant	Conditional Stability	Amount of excess chelate (ng ml ⁻¹)	Kinetic stability (dissociation half-life at pH 7.4)
Gadoversetamide, Gd-DTPA-BMEA (OligMark, Tyco, St. Louis, MO)	Non-ionic linear	15.5	15	28.4	Not available
Gadobutamide, Gd-DTPA-BMEA (Omniscan, GE, Waukesha, WI)	Non-ionic linear	16.9	16.9	12	25 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 (Magnevist, Schering, Berlin, Germany)	ionic linear	22.1	18.1	0.6	10 min
Gadobenate, Gd-BOPTA (Munipance, 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

Ultrasound



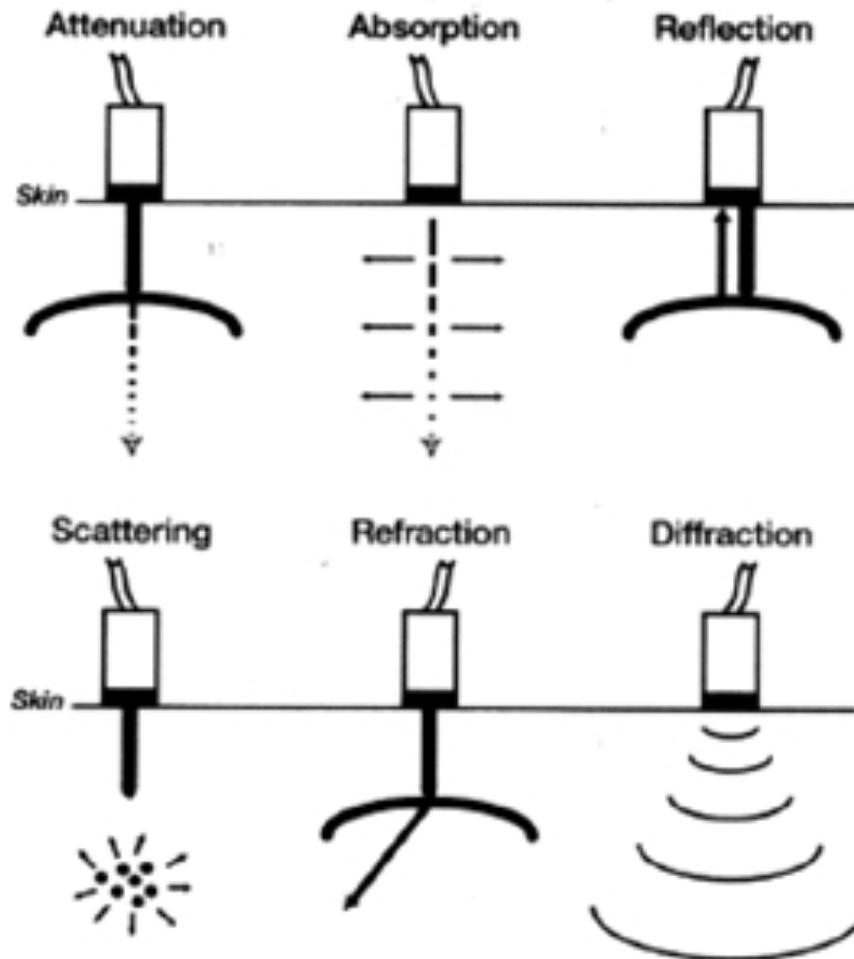
Ultrasound basics

US basics



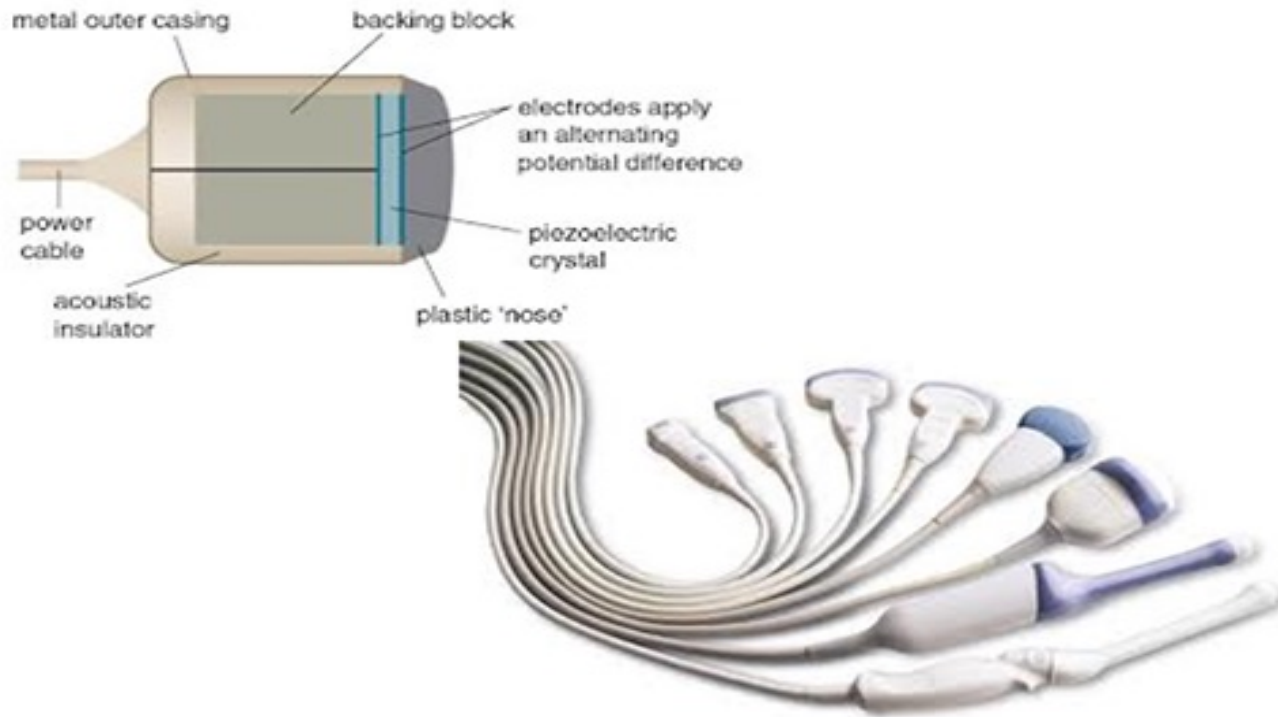
Imaging dependent on the speed of sound
In tissue

Fate of sound waves in body

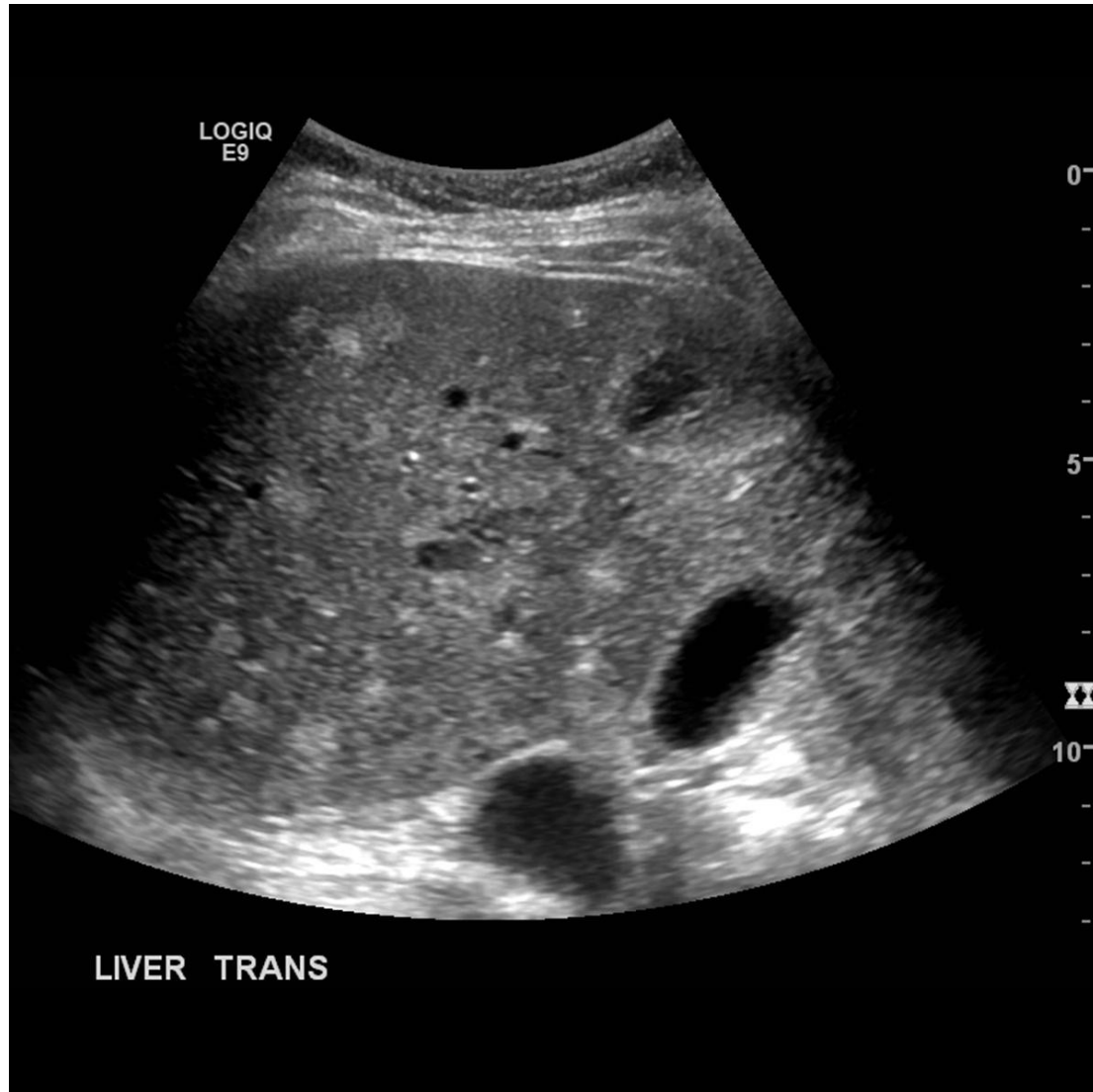


Ultrasound probes

US Probes



Liver metastases

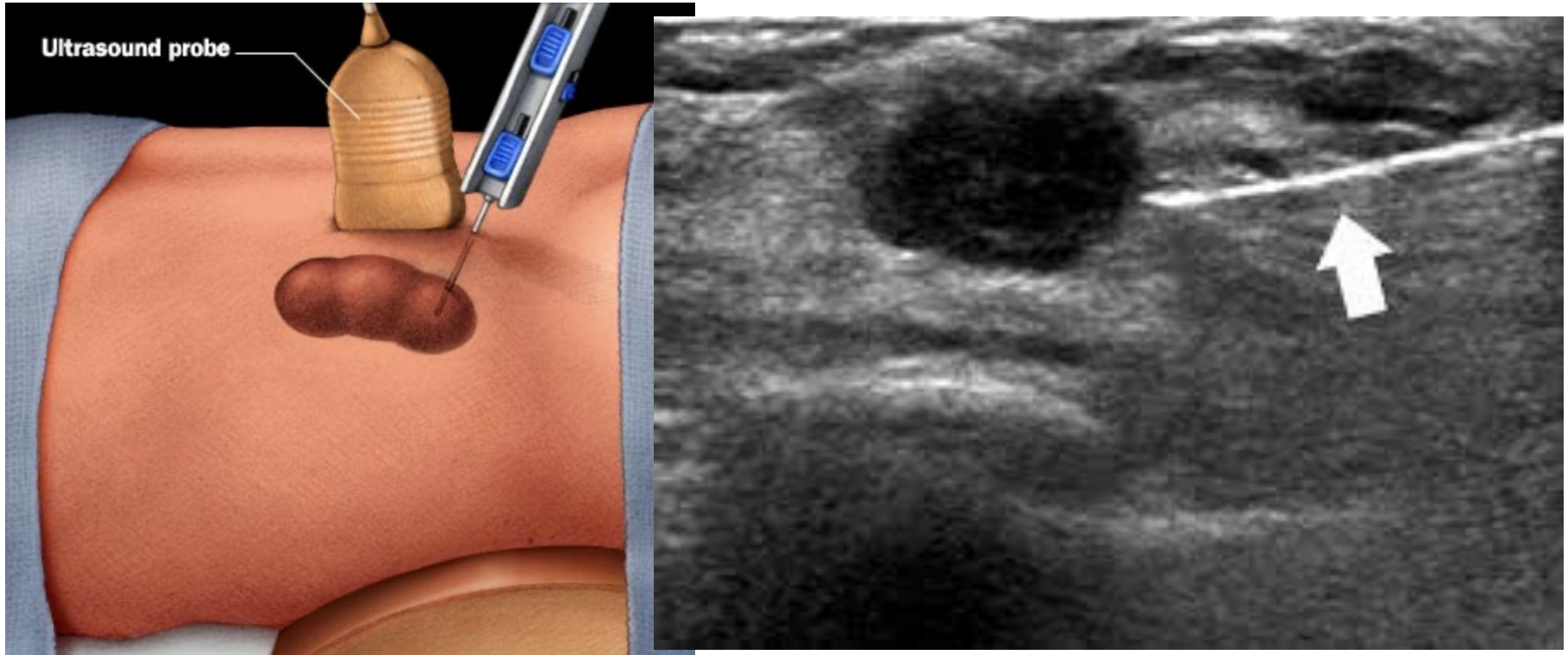


Ultrasound devices

Evolution of US devices



US guided biopsy-real time



US advantages

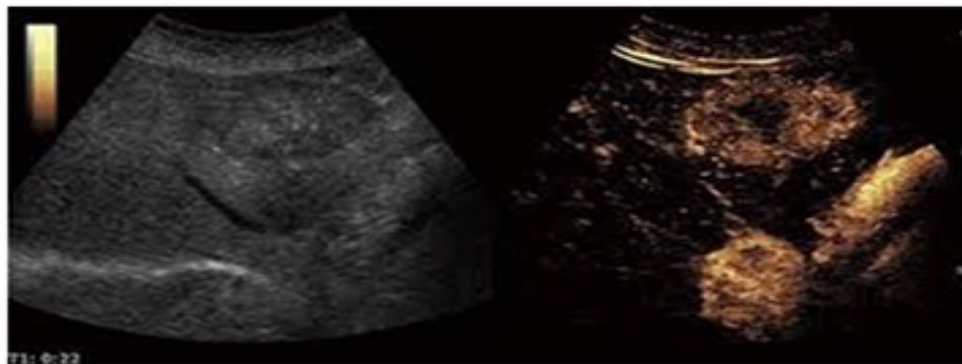
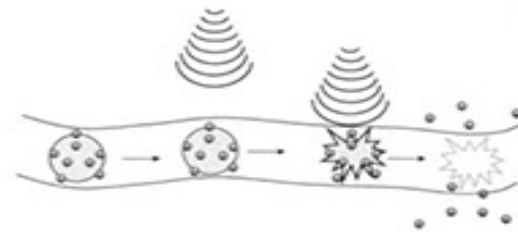
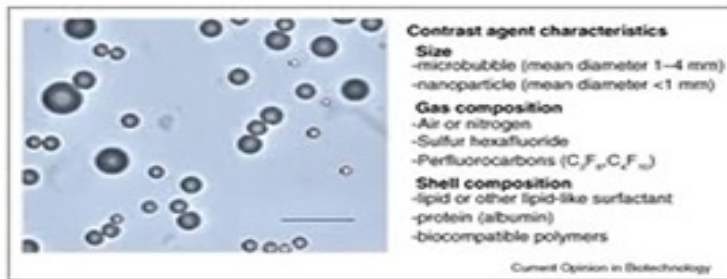
- No radiation
- Real time
- Inexpensive
- Quick, little prep
- No injection

US disadvantages

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

Microbubble contrast

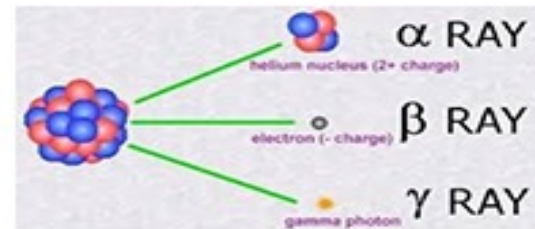
US Microbubble contrast



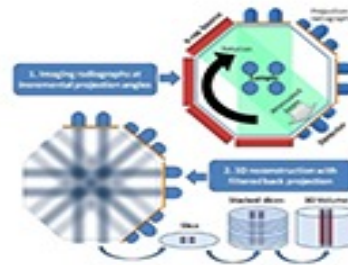
SPECT

Single Photon Emission Computed Tomography-SPECT

- Single Photon Emission



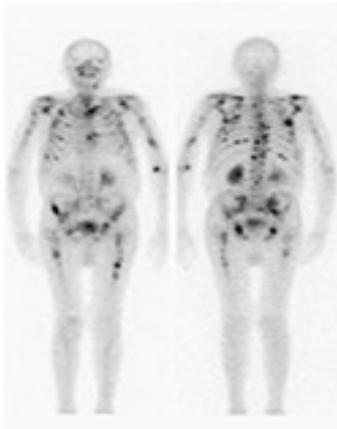
- Computed Tomography



SPECT imaging

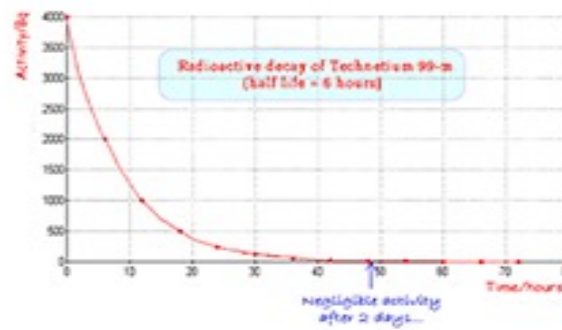
SPECT Imaging

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

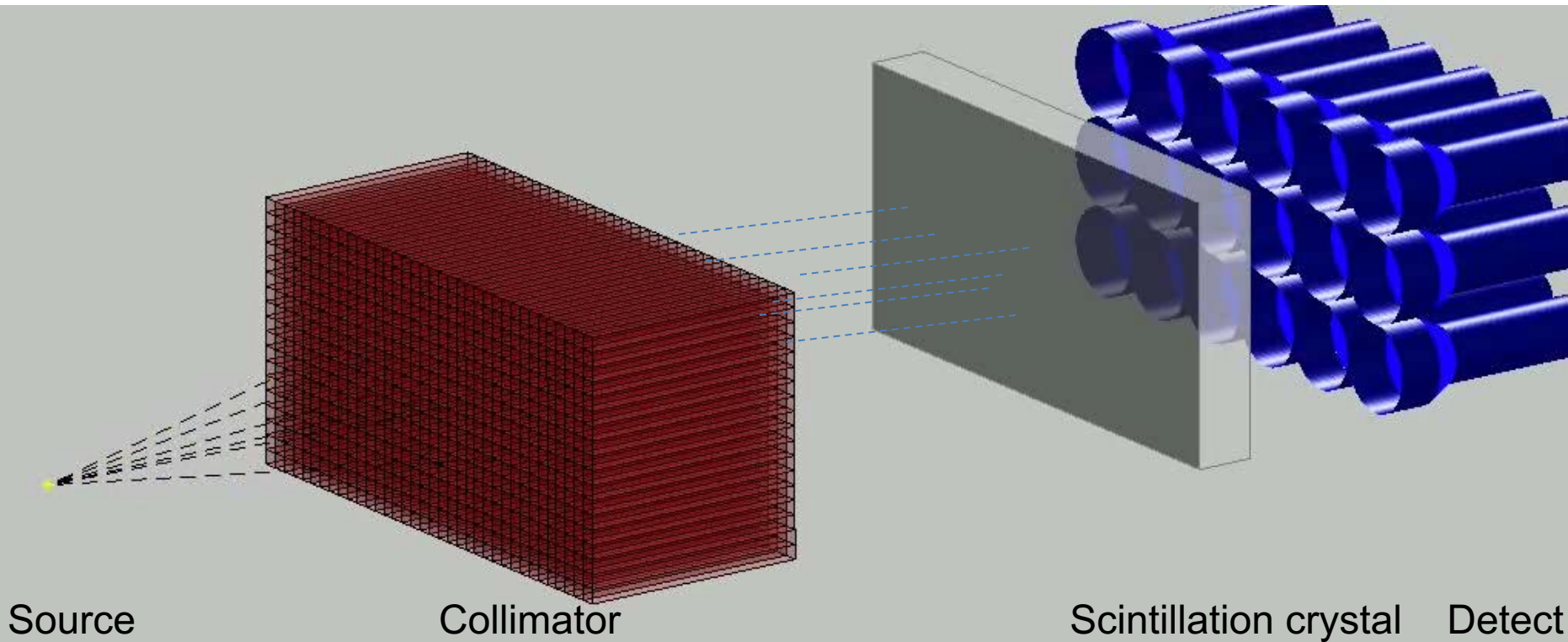


The bone scan:

^{99m}Techneium-methyl diphosphonate

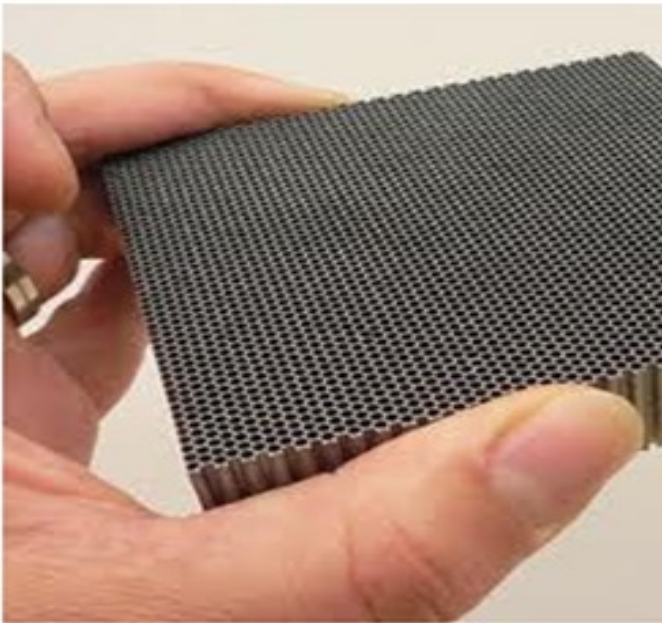


SPECT detectors



Collimation

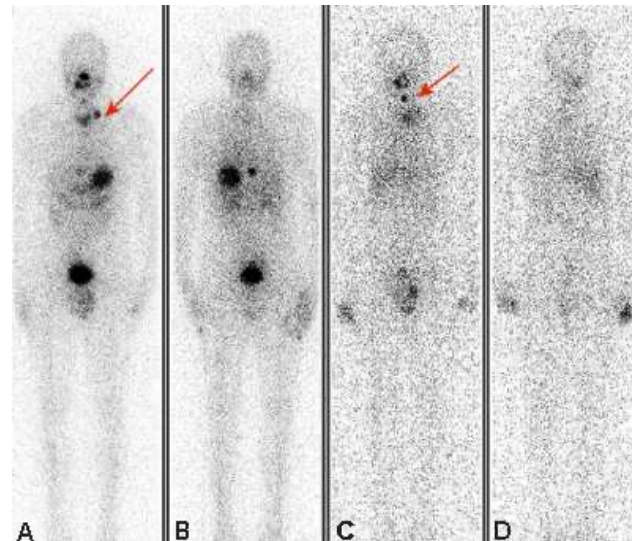
Collimation cont'd



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

SPECT agents for cancer

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



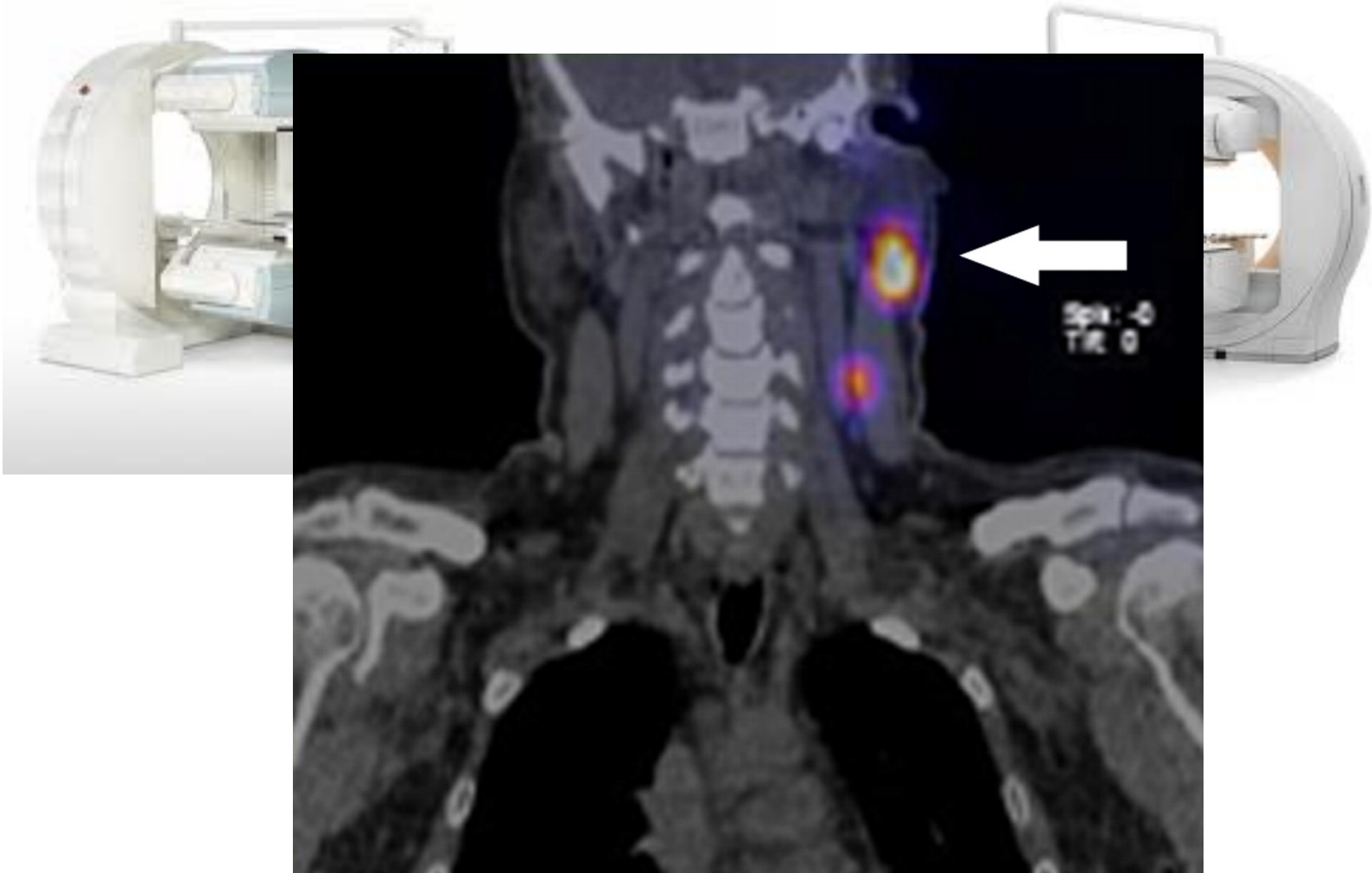
SPECT

SPECT Advantages/Disadvantages

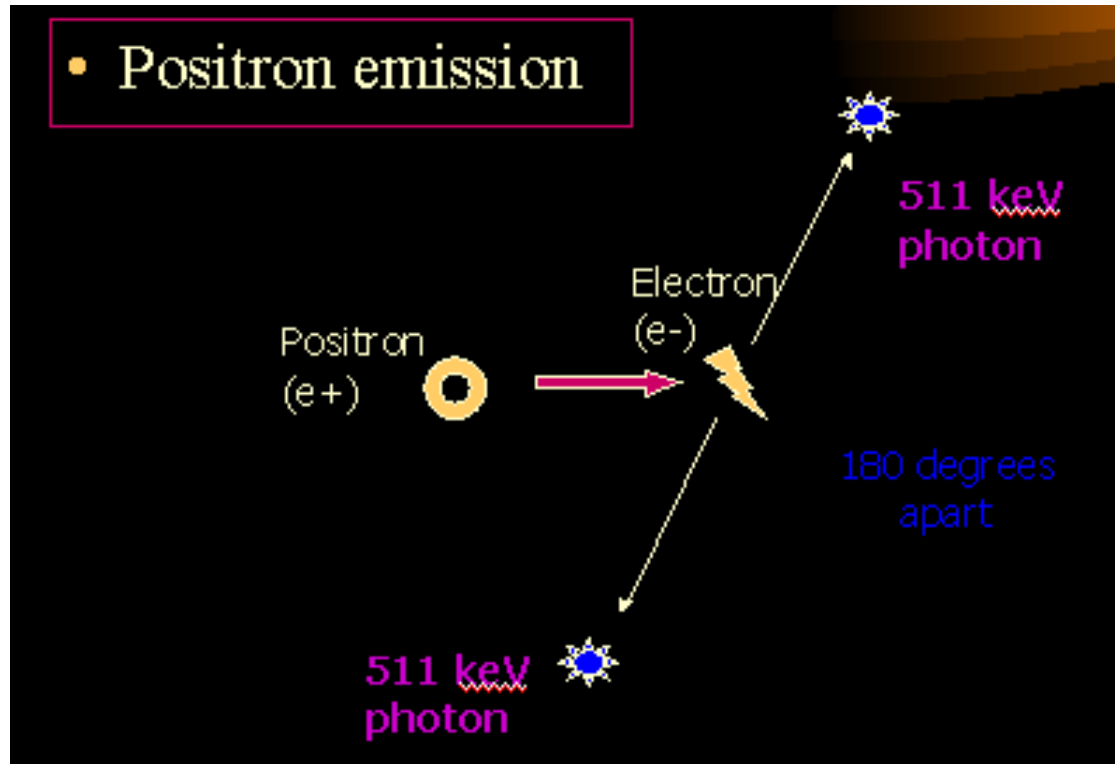
- Relatively inexpensive
- Broad experience
- Disadvantages
 - Radiation exposure
 - Preparation of imaging agent
 - Nuclear Regulatory
 - Scanning is slow, low resolution



Hybrid Imaging

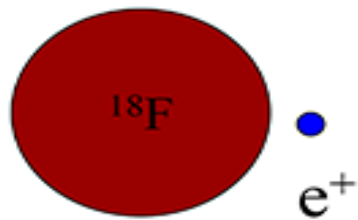


Positron Emission Tomography

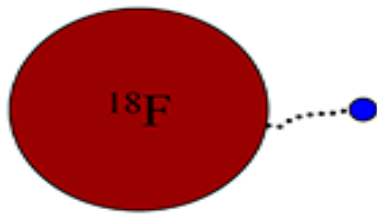


PET1

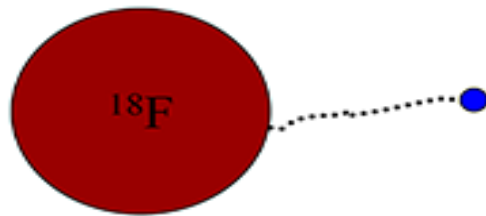
Positron Emission Tomography



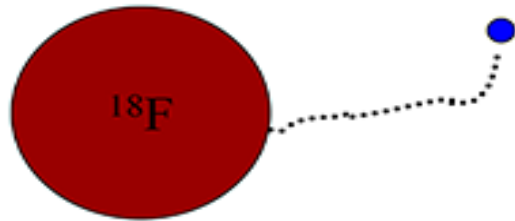
PET2



PET3



PET4



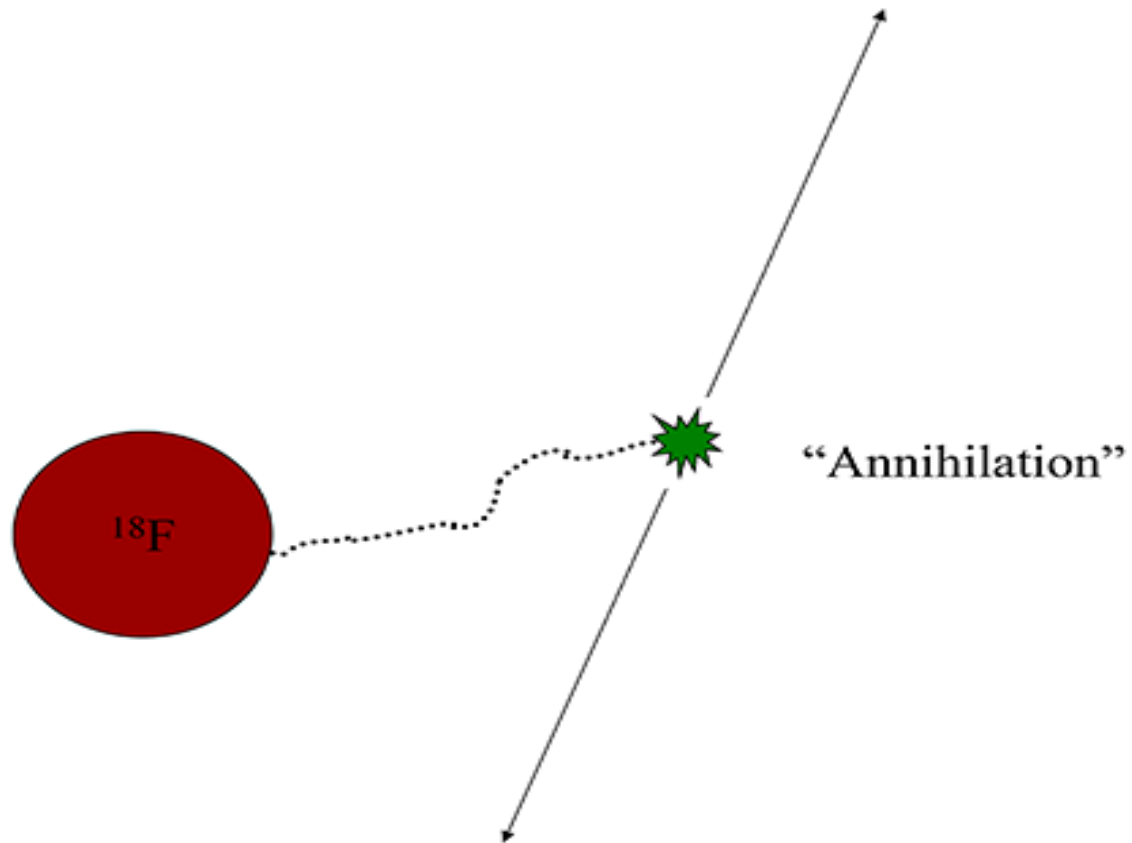
PET5



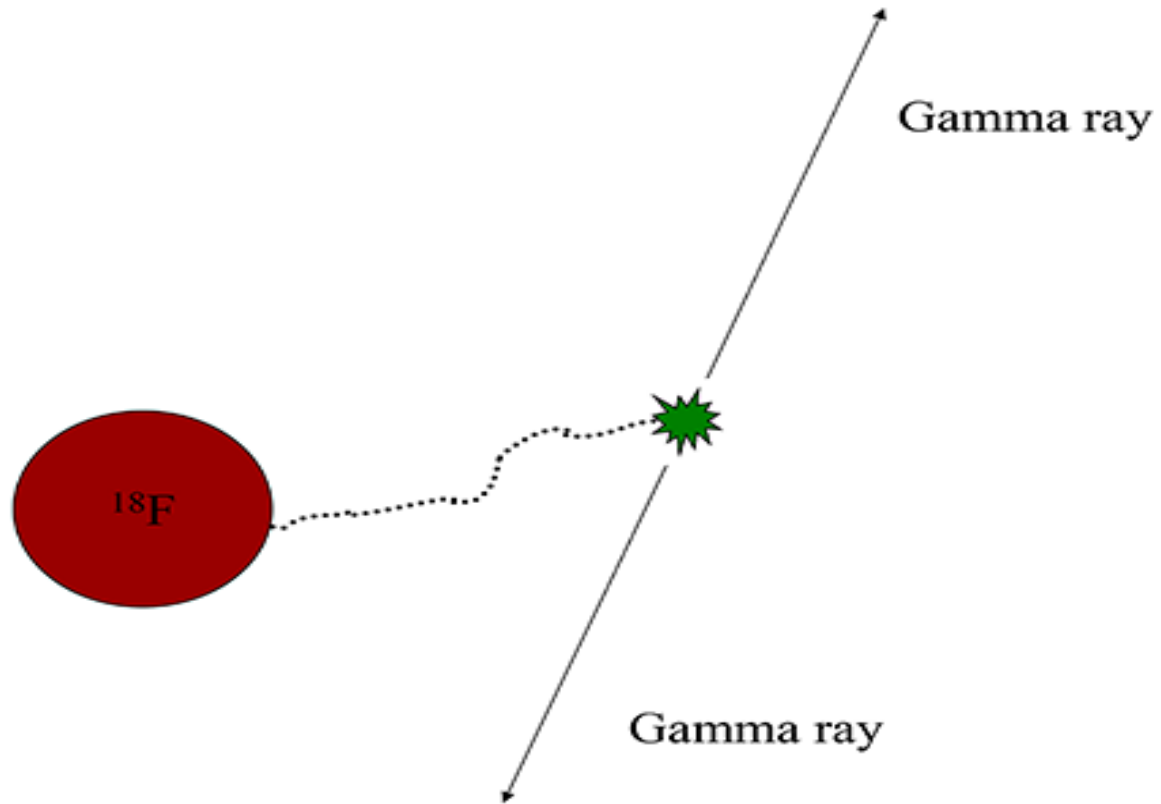
PET6



PET7

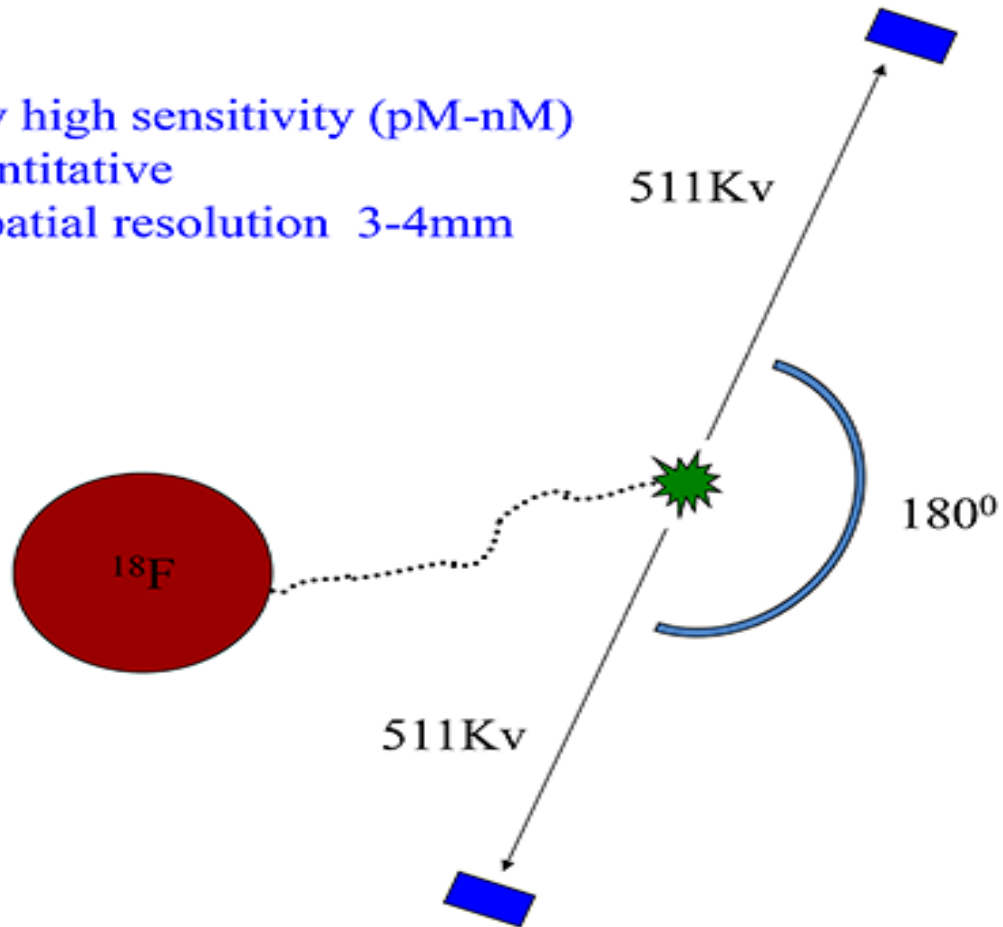


PET8



PET9

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



F-18 Deoxyglucose

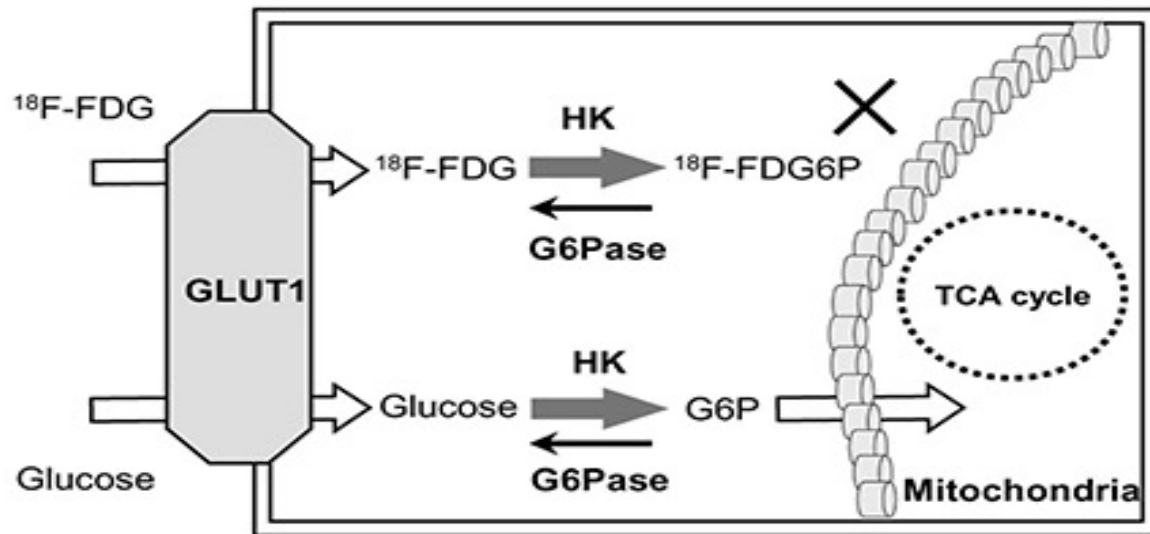
F-18 Deoxyglucose



Otto Warburg

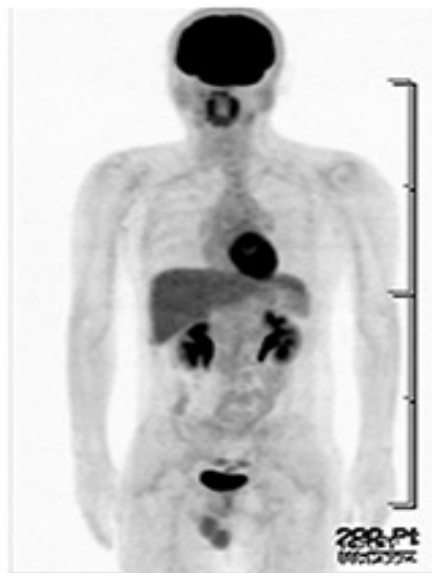


Lou Sokoloff

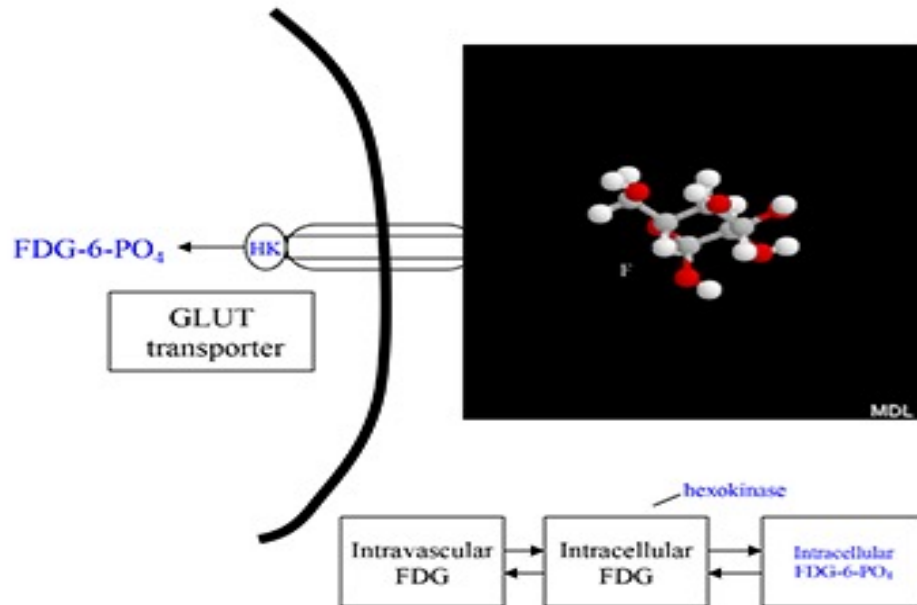


PET imaging

^{18}F FDG PET Imaging

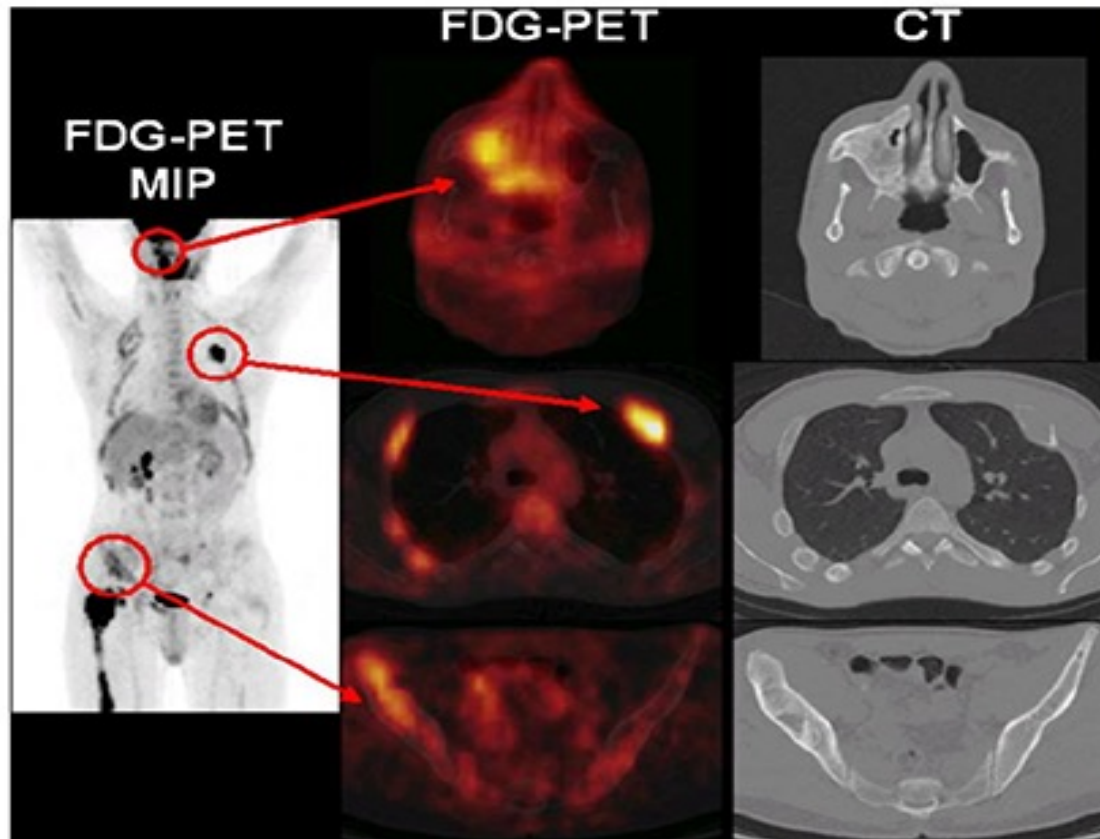
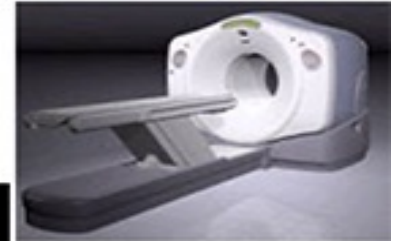


E PET MIP

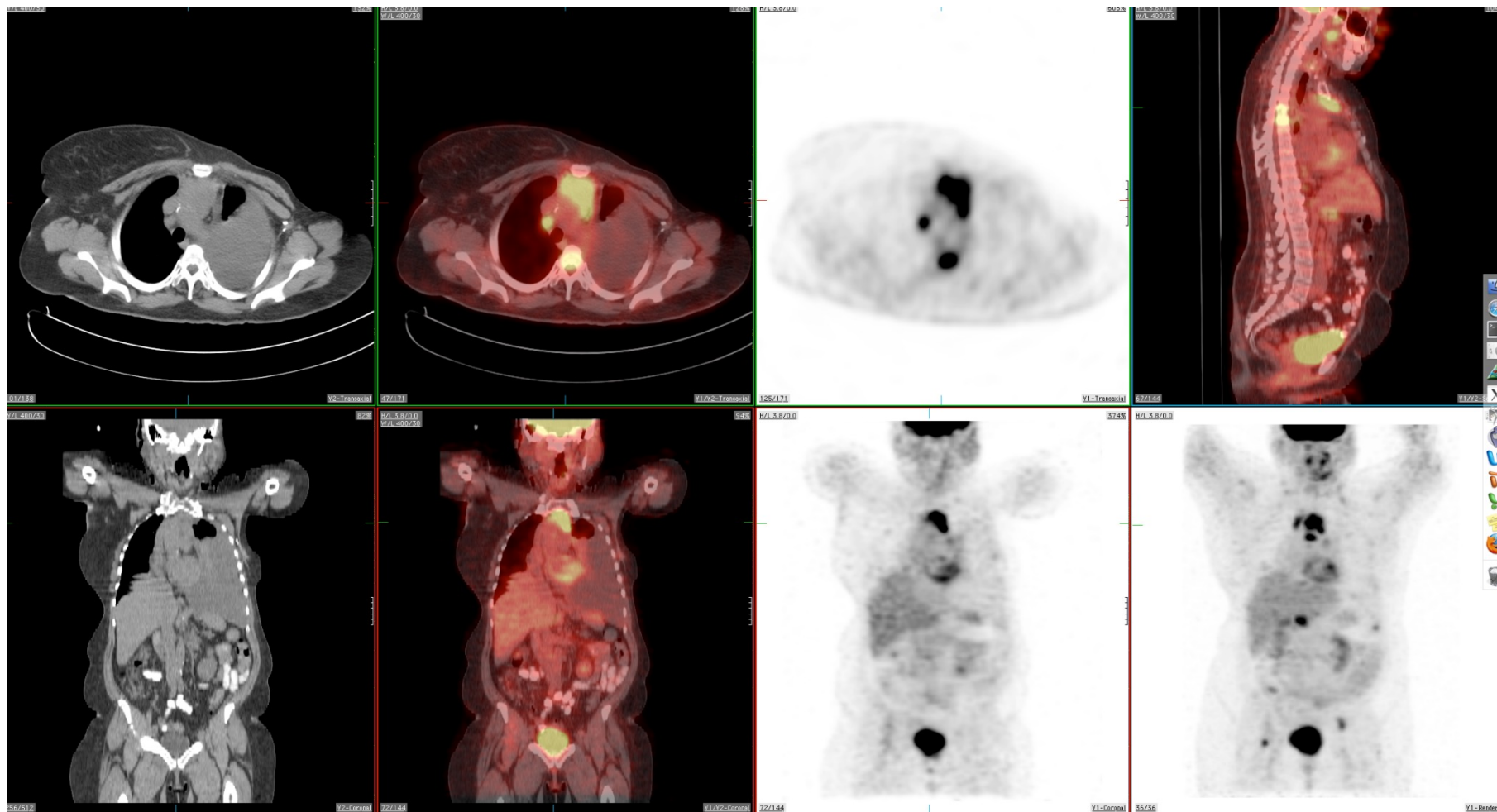


PET scanners

PET-CT scanners



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

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

Presentation

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

Resolution

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

Sensitivity

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

Cost

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

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>

pchoyke@nih.gov