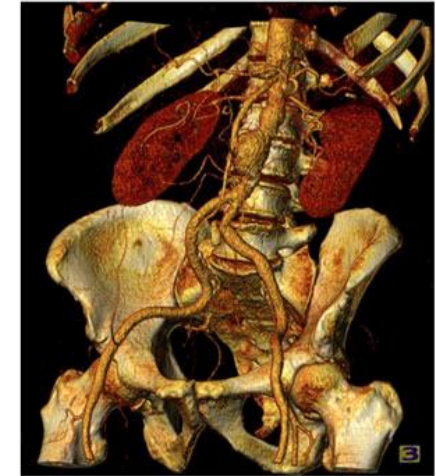
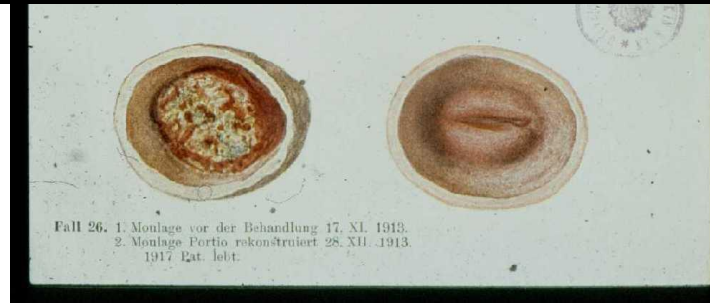


Image Guided Radiotherapy

Richard Pötter, MD,
supported through Markus Stock PhD, Med Austron
Wolfgang Marik MD, Department of Radiology,
Maximilian P Schmid MD, Department of Radiotherapy
Dietmar Georg, PhD, Medical Physics,
Peter Kuess, PhD, Medical Physics
Medical University of Vienna

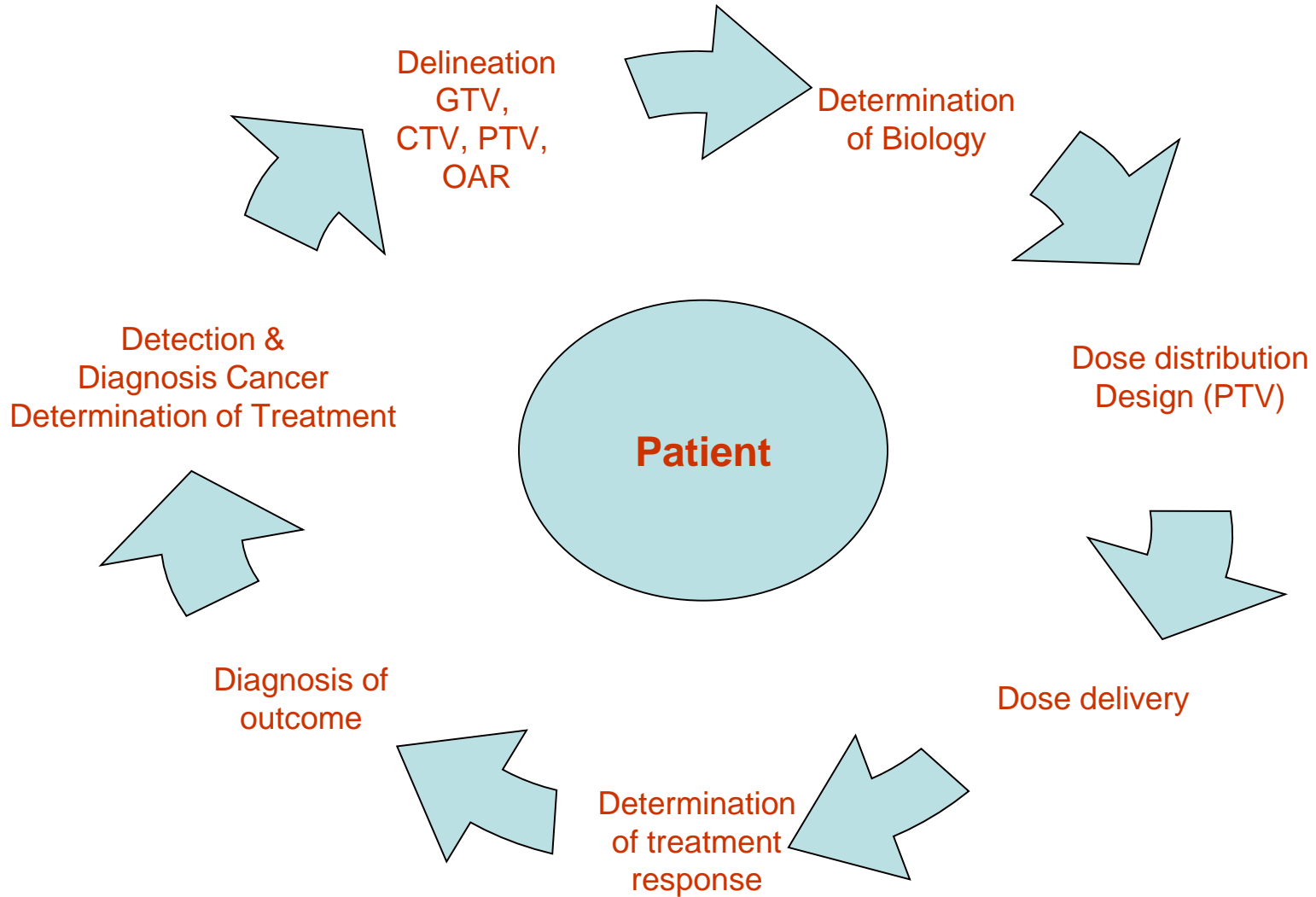
**Image Guided RadioTherapy (IGRT):
a comprehensive view (7Ds)**

What is medical imaging?



- Techniques and processes used to create images of the human body for clinical purposes
- Radiography, Fluoroscopy, CT, Ultrasound, MRI, PET CT, Endoscopy, Cartoons...
- How is medical imaging used in Radiotherapy?

Imaging-Cycle



(D1) Detection, Diagnosis, Determination of treatment

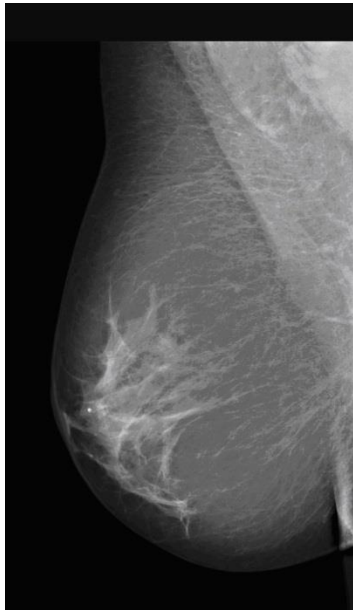
- improvements in (cancer-specific) survival -

- Screening programs (Mammography, endoscopy, tests)
- Diagnosis at earlier stage → more effective treatment options (cervix, breast, colorectal, prostate, stomach, oesophagus...)
- More appropriate stage (risk) assessment (CT/MRI)
more appropriate treatment strategy allocation
- More appropriate spread assessment (e.g. PET CT)
Decision: local (where), regional, systemic approach (e.g. lung)

Common screening programmes

Breast Cancer

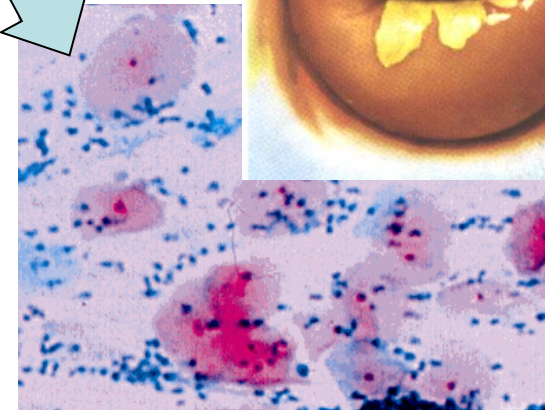
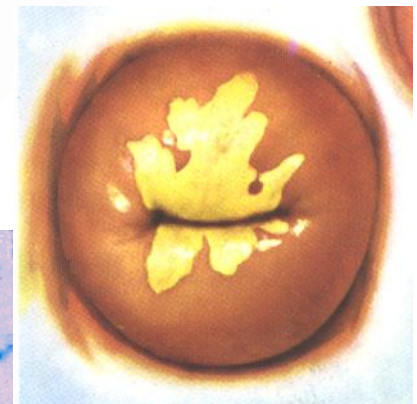
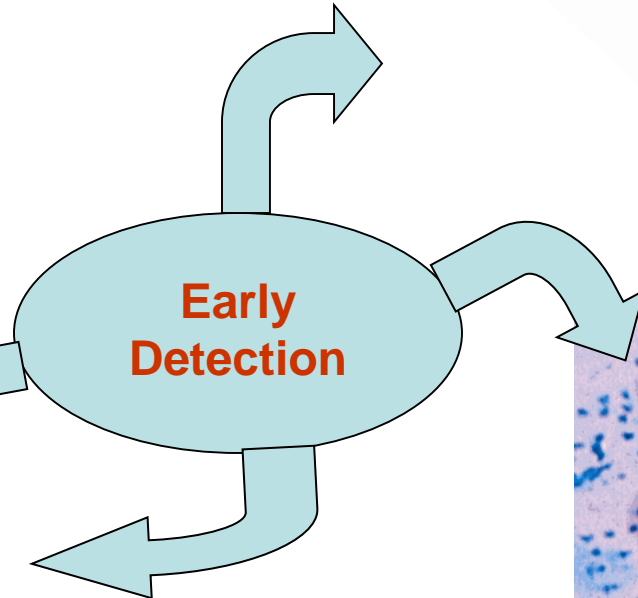
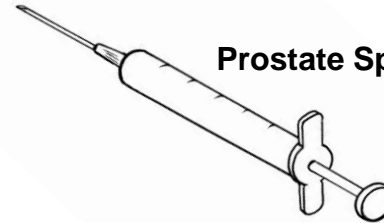
mammography



Tests (blood,...)

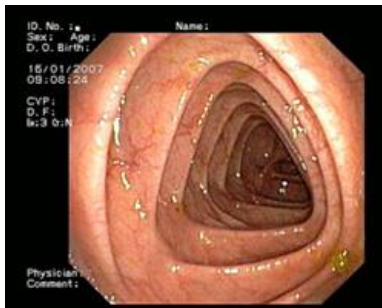
Prostate Specific Antigen (PSA)

Prostate Cancer



Cytology

Cervix Cancer

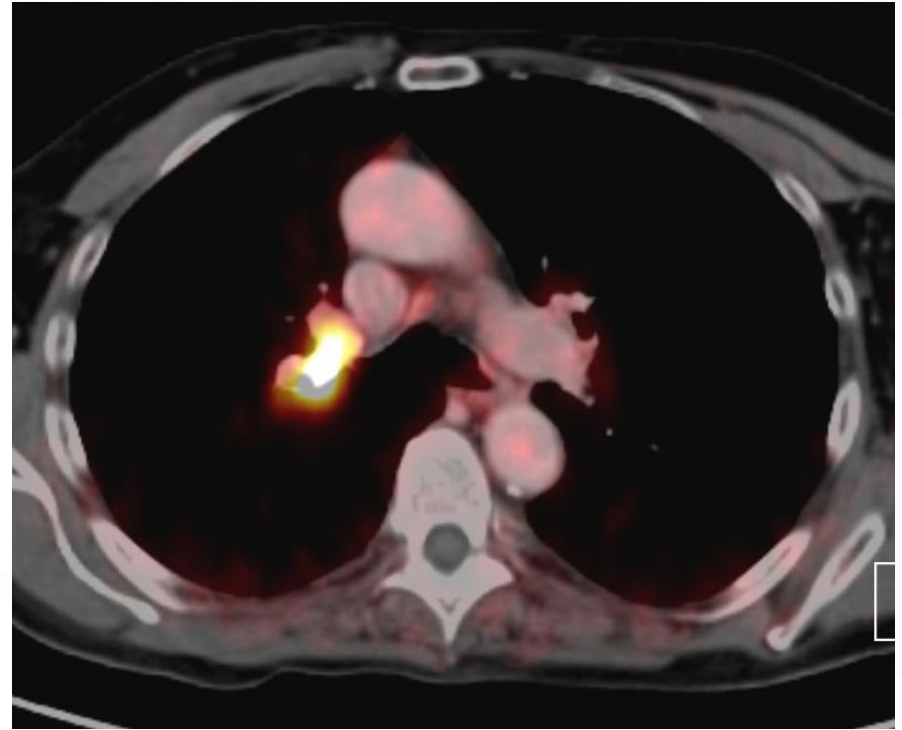
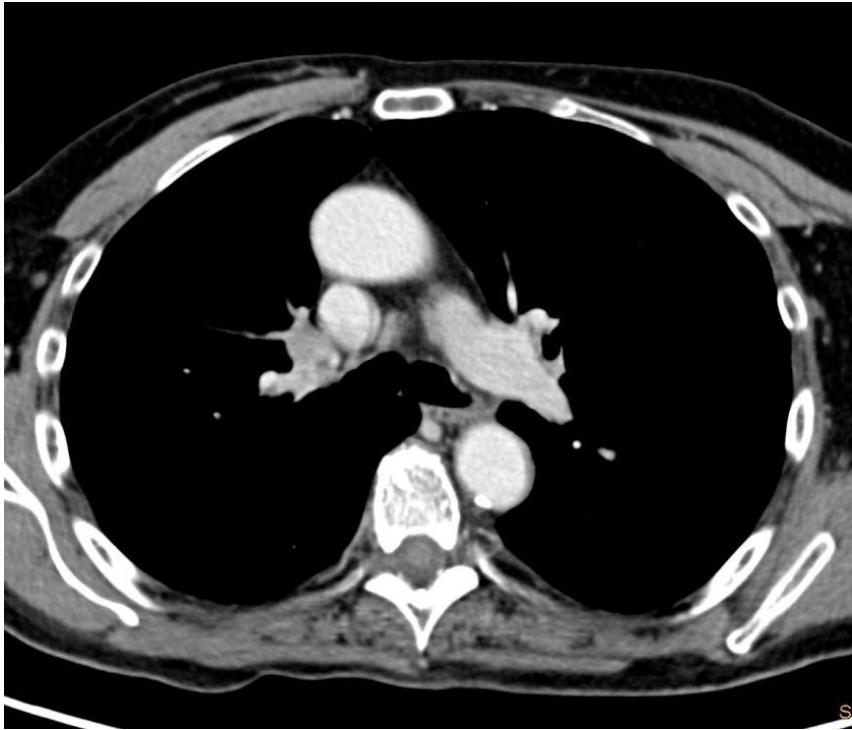


Colonoscopy

Colorectal Cancer

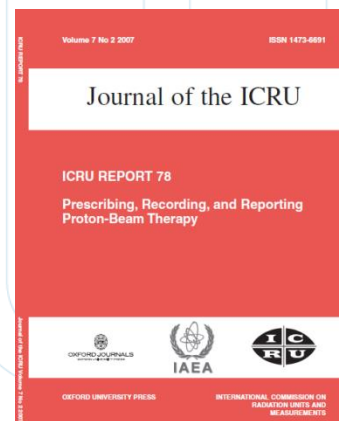
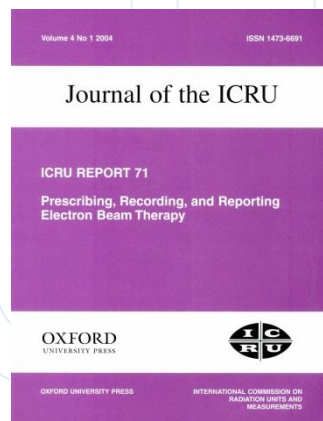
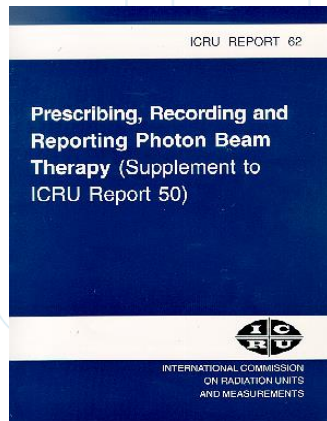
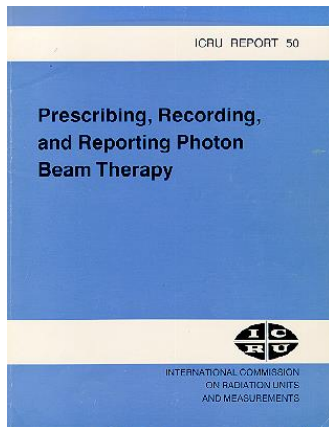
Lung Tumor

CT, PET CT



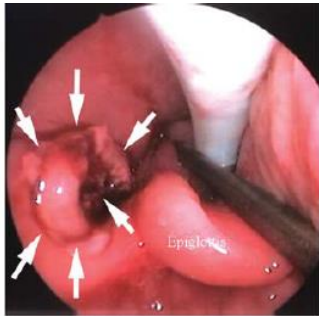
Tumor/Target volume and dose volume concepts in Radiation Oncology

- The **International Commission on Radiation Units and Measurements (ICRU (since 1927))** has as its principal objective the development of internationally acceptable recommendations
 - Quantities and units of radiation and radioactivity
 - **Procedures** suitable for the measurement and application of these quantities in clinical radiation oncology and radiobiology



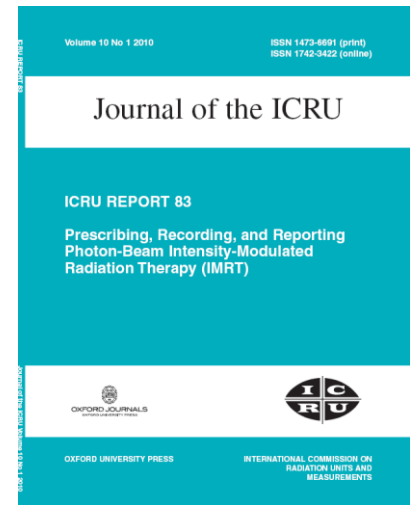
ICRU defines a common language for clinical practice and for medical and scientific communication in Radiation Oncology

Gross Tumor Volume (GTV)



Laryngeal cancer view from a laryngoscope

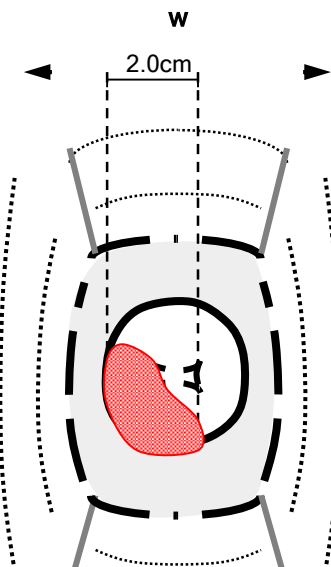
- the gross palpable or visible / demonstrable extent and location of malignant growth.
- based on information from a combination of diagnostic modalities
 - clinical examination, endoscopy (light imaging)
 - X-Ray, CT, MRI, ultrasound, PET CT, etc.,
 - Histology after biopsy
- **TIME AND IMAGING MODALITY ARE IMPORTANT!!!**



At diagnosis

IIb

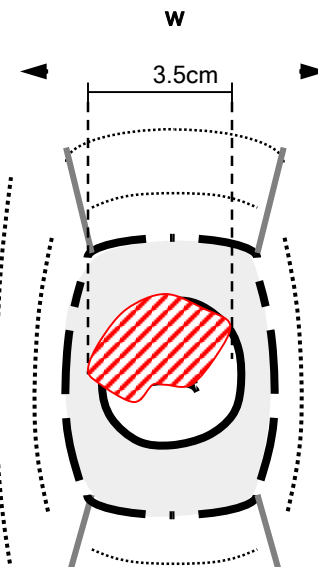
$w = 2.0 \text{ cm}$
 $h = 2.0 \text{ cm}$
 $t = 1.5 \text{ cm}$



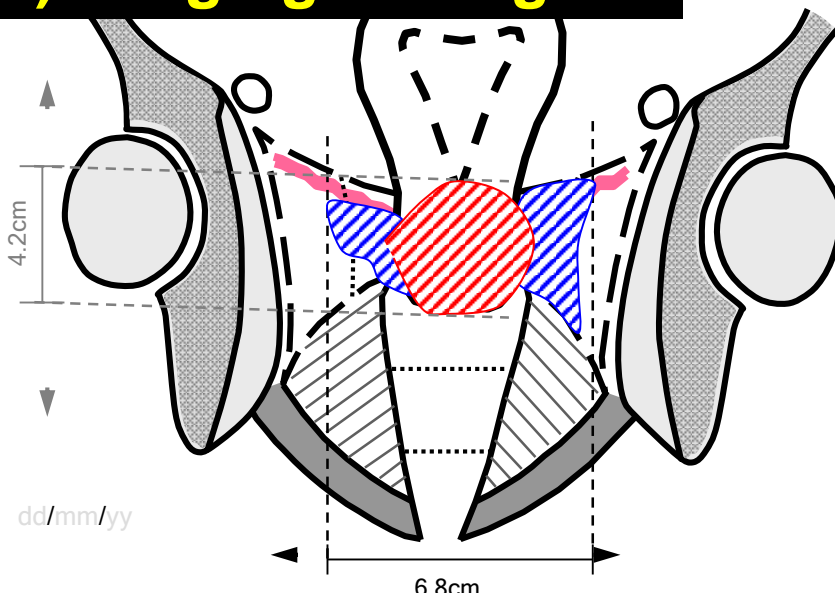
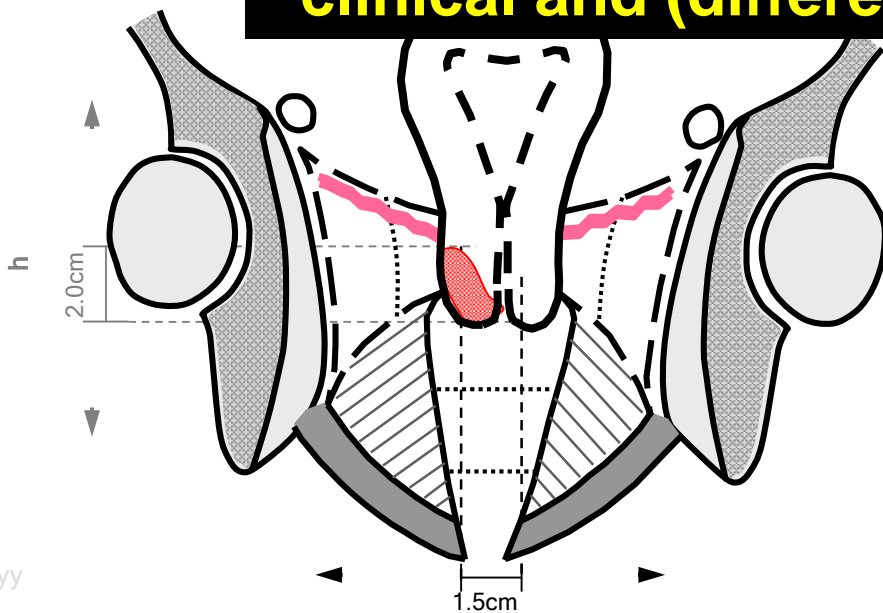
At diagnosis

IIIb

$w = 6.8 \text{ cm}$
 $h = 4.2 \text{ cm}$
 $t = 4.5 \text{ cm}$



**Cartoon drawings for comprehensive view
clinical and (different) imaging findings**

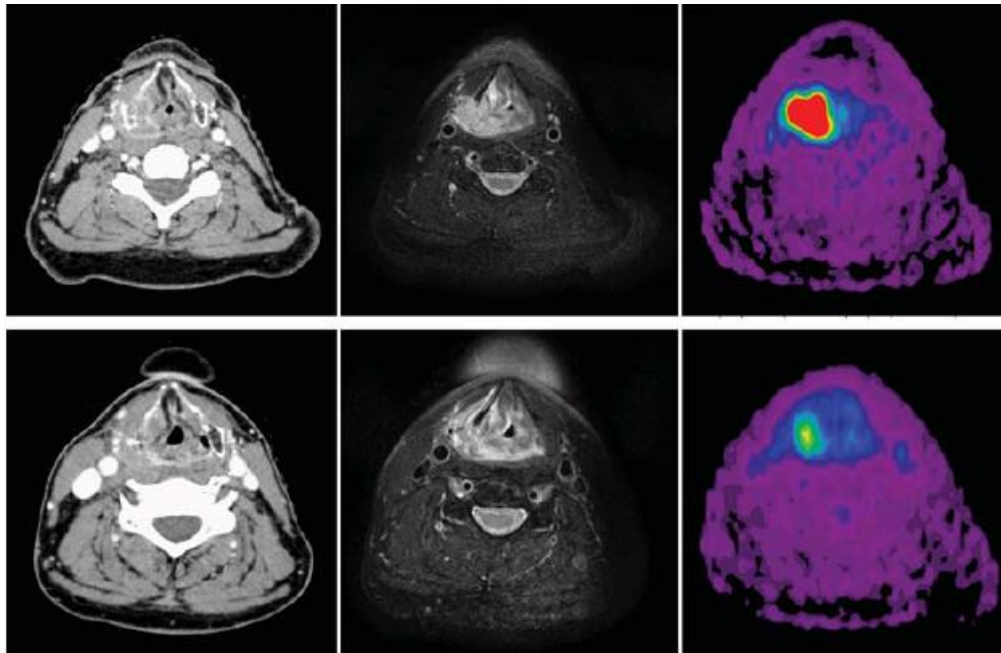


dd/mm/yy

dd/mm/yy

Gross Tumor Volume (GTV)

- Comparison among various modalities for the definition of the primary head-and-neck tumor GTV.
- Upper panel: GTV imaged prior to any treatment
 - contrast-enhanced CT: GTV-T: volume of 25.8 ml.
 - fat-saturated T2-weighted MRI: GTV-T: volume of 28.5 ml.
 - FDG-PET: GTV-T: volume of 22.2 ml.

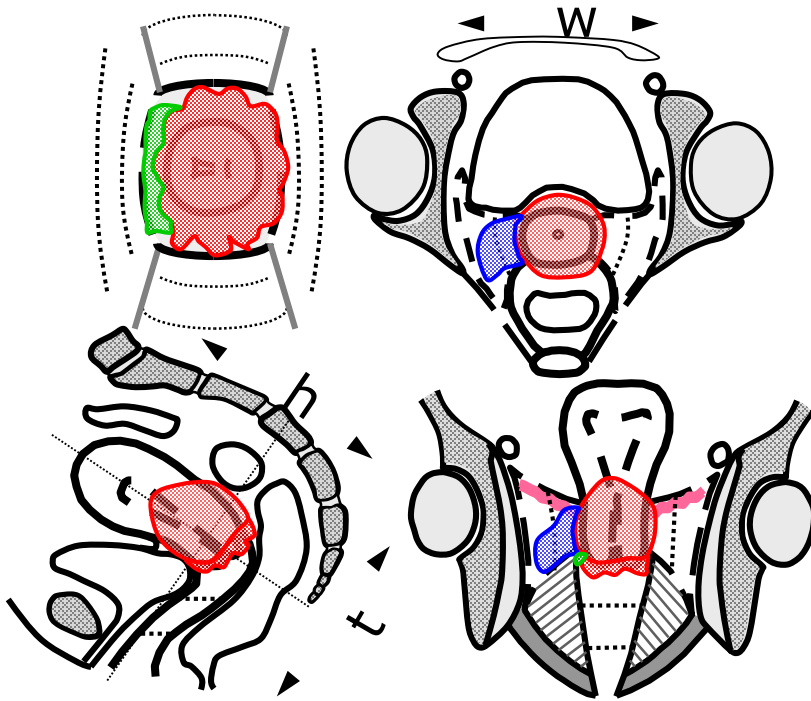


GTV during treatment

Lower panel: GTV imaged after an absorbed dose of 20 Gy.

- GTV-T (CT, 20 Gy): volume of 16.3 ml.
- GTV-T (MRI T2, fat sat, 20 Gy): volume of 19.8 ml.
- GTV-T (FDG-PET, 20 Gy): volume of 12.5 ml.

Findings at time of diagnosis



Findings at time of brachytherapy

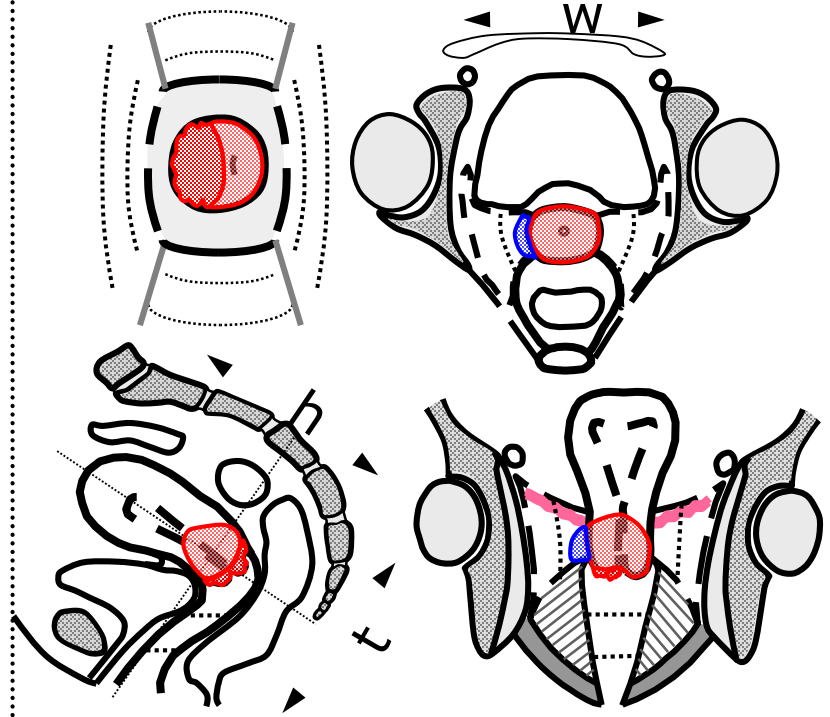
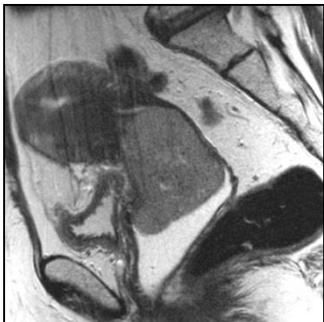
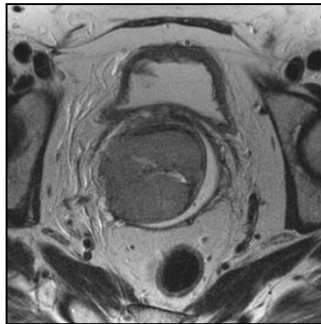


Fig.5.1:
GTV_{init}
and
GTV_{res}

in
cervix
cancer

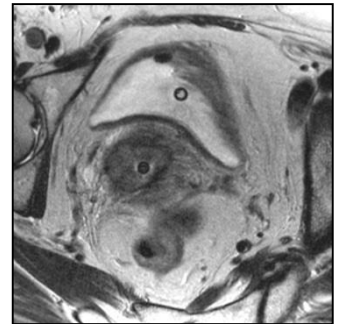
Dimensions (cm):

Width: 7
Thickness: >5
Height: >5
Vaginal inv.: 0.5
(right fornix)

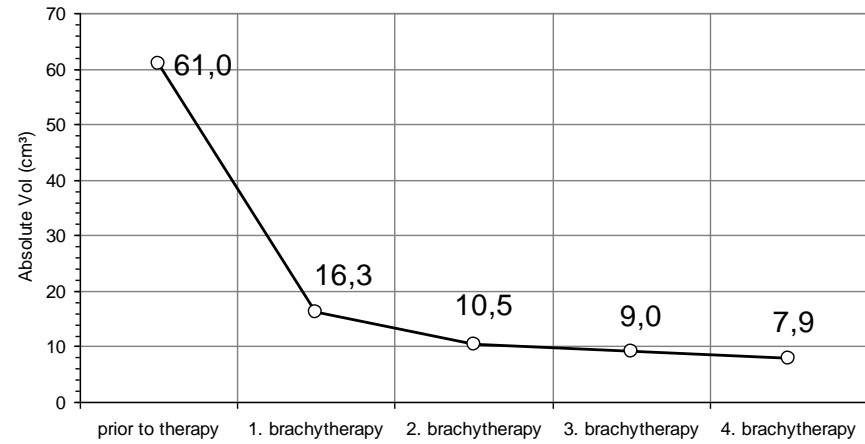
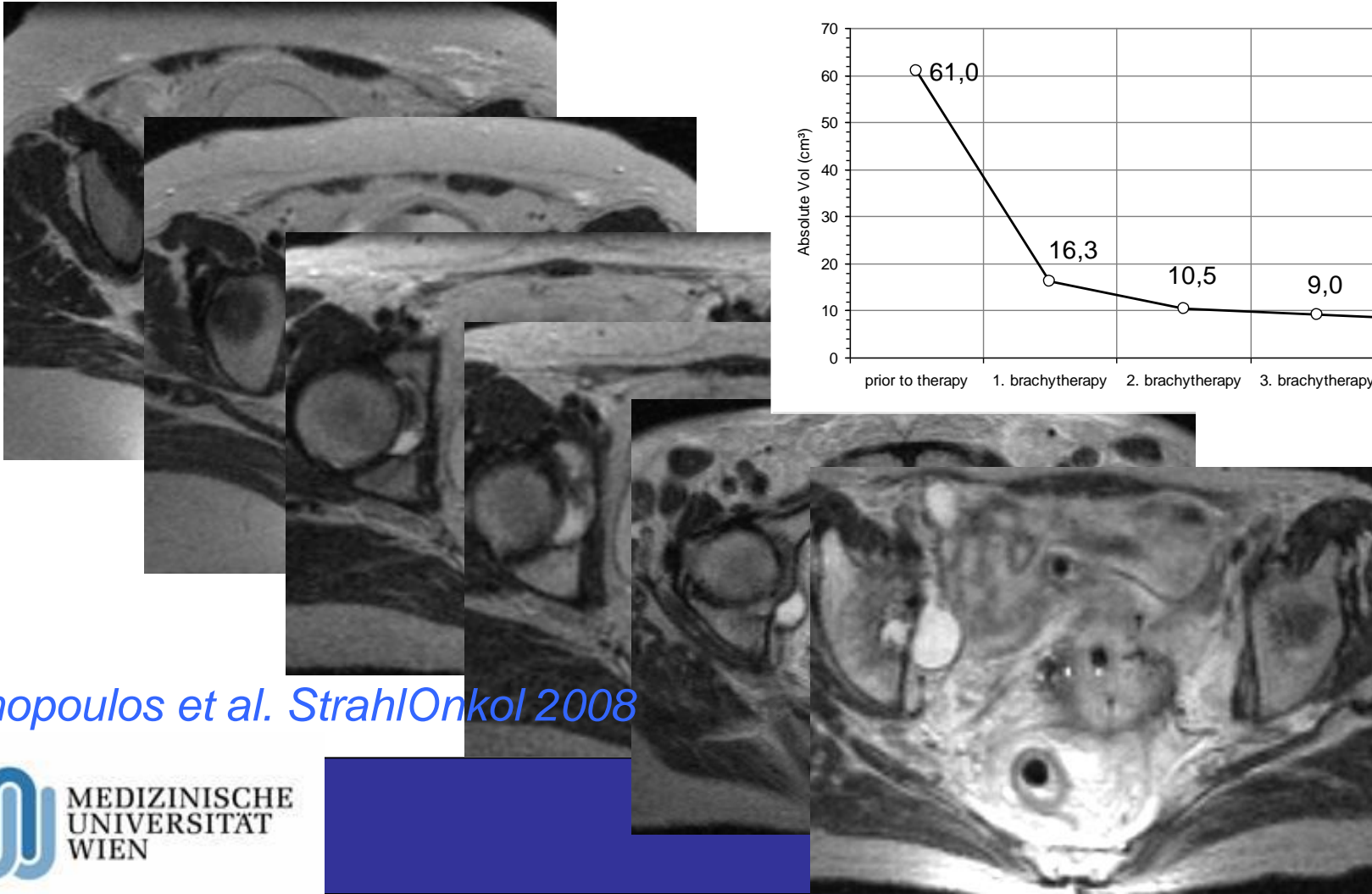


Dimensions (cm):

Width: 3.5
Thickness: 2
Height: 2
Vaginal inv.: 0

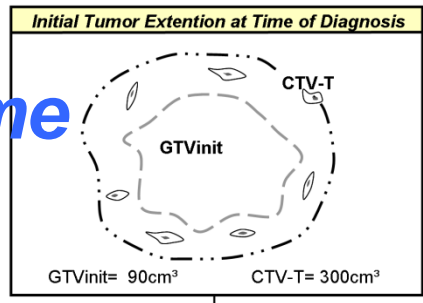


MRI: Initial tumour extension (3D RT) pattern of response (4D RT) for adaptive MRI based planning



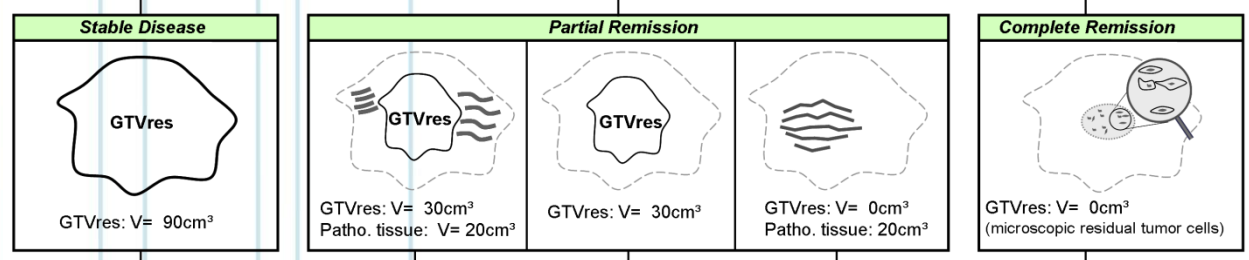
Dimopoulos et al. StrahlOnkol 2008

The challenge of change in tumour volume and tumour configuration during treatment

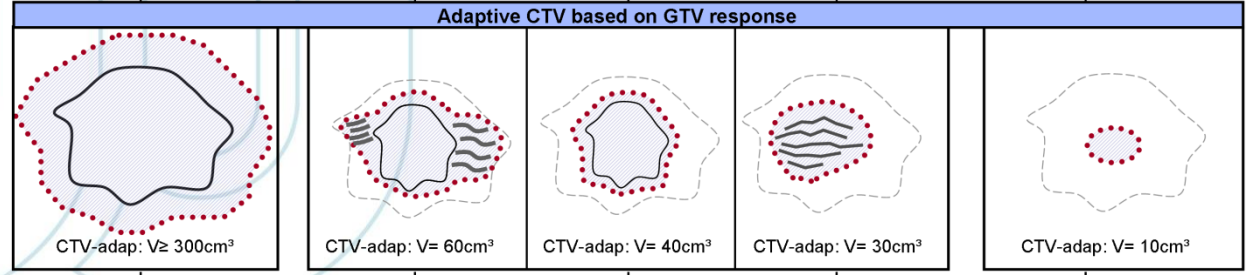


TREATMENT

Patterns of GTV Response



various patterns of



ADAPTIVE TREATMENT

 GTVinit
 CTV-T
 GTVres
 CTVadap
 Pathologic tissue

Various patterns
GTV response

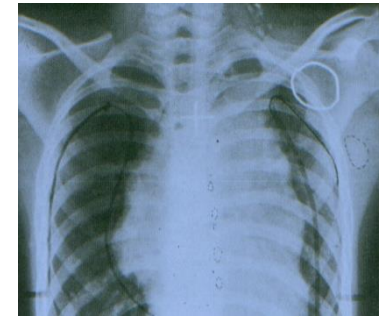
Corresponding
adaptive CTVs

ICRU/GEC ESTRO
Report 89
Fig 5.3

(D2) Delineation of GTV, CTV, PTV

+ organs at risk

- from planar imaging („2D“)
 - skin cancer (basalcelleptheliom)
 - X-ray Sim, individual blocks
 - Hodgkin´s Disease (2D-4D)
 - Boost Head and neck (clinical)(2D-4D)

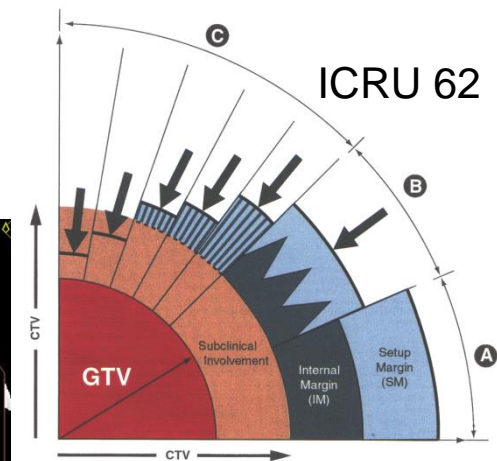
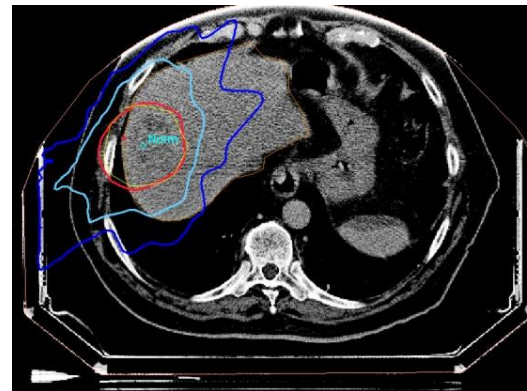


- to volumetric imaging („3D“)

- CT (MRI, US) based
- GTV, CTV, PTV
- ICRU 50&62

73&78&83&89

definitions

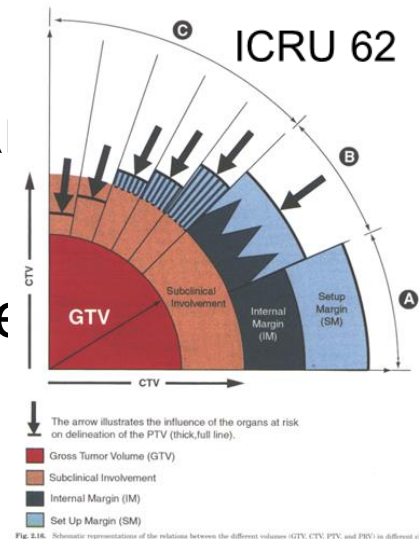


The arrow illustrates the influence of the organs at risk on delineation of the PTV (thick, full line).

- Gross Tumor Volume (GTV)
- Subclinical Involvement
- Internal Margin (IM)
- Set Up Margin (SM)

Volume definition (ICRU language)

- The process of determining the volume for the treatment of a malignant disease consists of **several distinct steps**.
- **Different volumes** may be defined, e.g. due to:
 - varying (assumed) concentrations of malignant
 - probable changes in the spatial relationship between volume and beam during the
 - movement of patient
 - possible inaccuracies in the treatment setup.
- **ICRU Reports define and describe several volumes:**
 - GTVs, targets and normal structures
 - to aid in the treatment planning process
 - to provide a basis for treatment comparisons



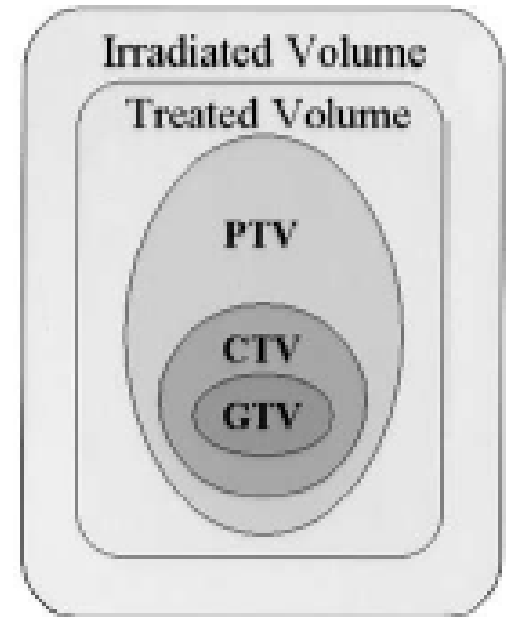
ICRU Report 50

- 1993 - 3D-CRT

GTV *visible tumor*

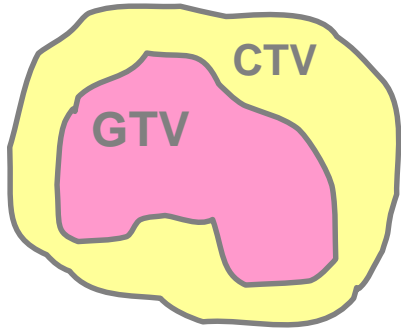
CTV = *GTV + microscopic spread (lymph node, perivascular, perineural)*

PTV = *CTV + geometric uncertainties (organ motion, tumor & patient movement, inaccuracies of beam & patient setup)*



- Appearance of new imaging modalities (MR, PET), technological progress (virtual simulation, MLC, IMRT), more information about target/organ movement lead to
 - Report 62 (Supplement to Report 50)

Clinical Target Volume (CTV)



- the tissue volume that contains a demonstrable GTV and/or sub-clinical microscopic malignant disease, which has to be eliminated.
- often includes an area directly surrounding the GTV that may contain microscopic disease and other areas considered to be at risk and require treatment.
- is an anatomical-clinical volume:

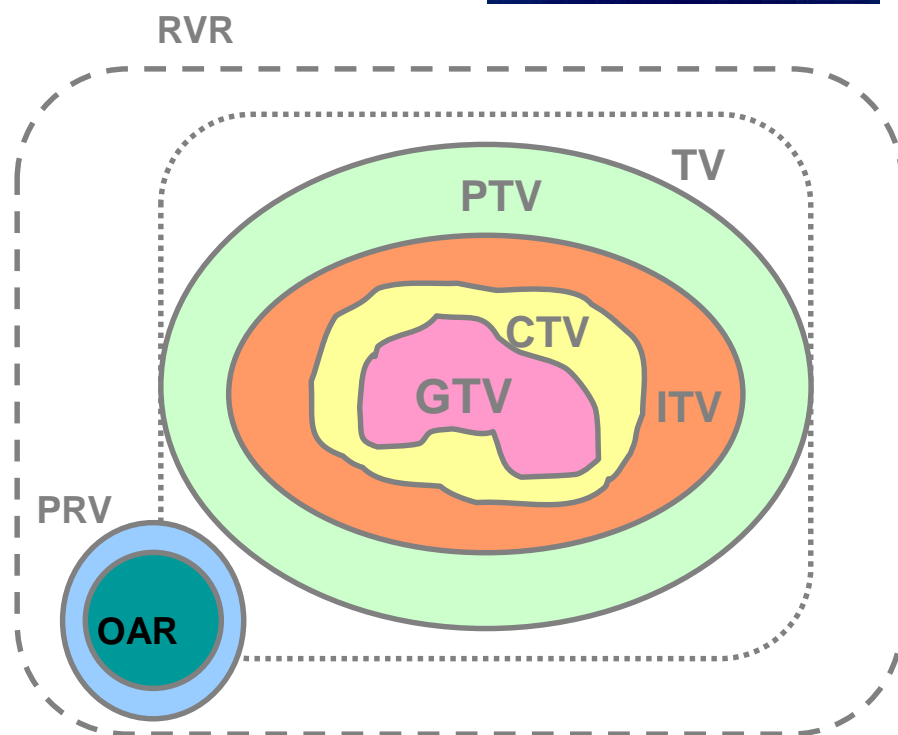
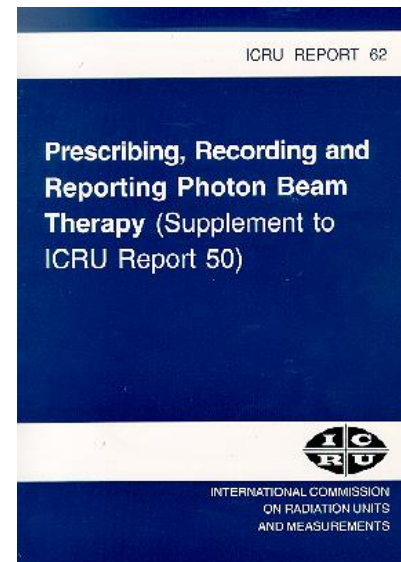
CTV-T (Tumor),

CTV-N (involved lymph Node)

CTV-M (Metastasis)

ICRU Report 62

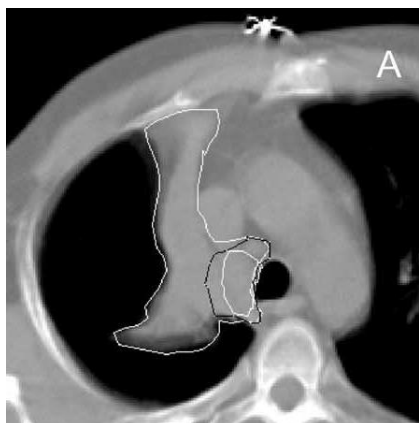
- Gross tumor volume: GTV
- Clinical target volume: CTV
- Organ at risk: OAR
- Planning target volume: PTV
- Internal target volume: ITV
- Treated volume or TV
 - Volume enclosed by specific isodose (D98%)
- Planning organ-at-risk volume: PRV
- Remaining volume at risk: RVR
 - Remaining volume at risk
 - (Patient – (CTV+OARs))



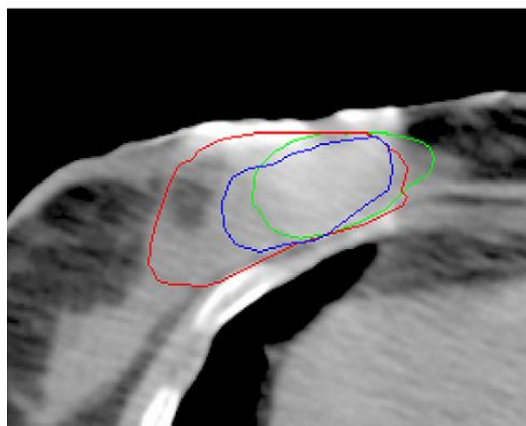
Volume definition

Target Volume definition

InTerobserver variability in delineation...

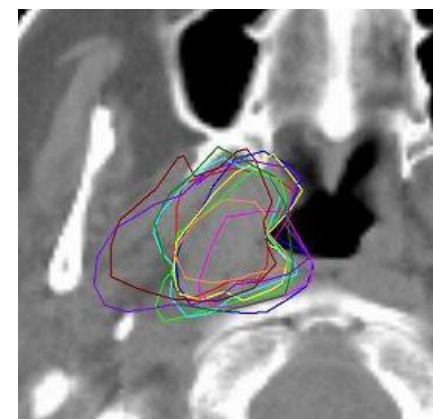
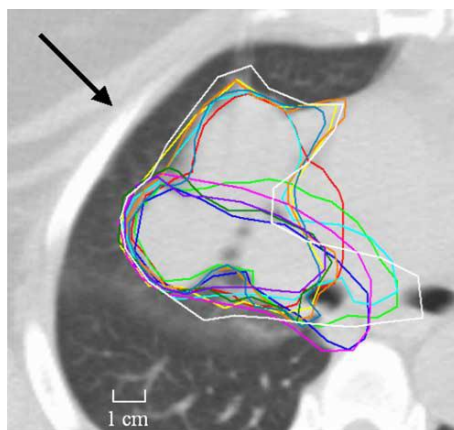


J. Van de Steene et al. / Radiotherapy and Oncology 62 (2002) 37-49



R. P. Petersen et al.

I. J. Radiation Oncology ● Biology ● Physics Volume 69, Number 1, 2007

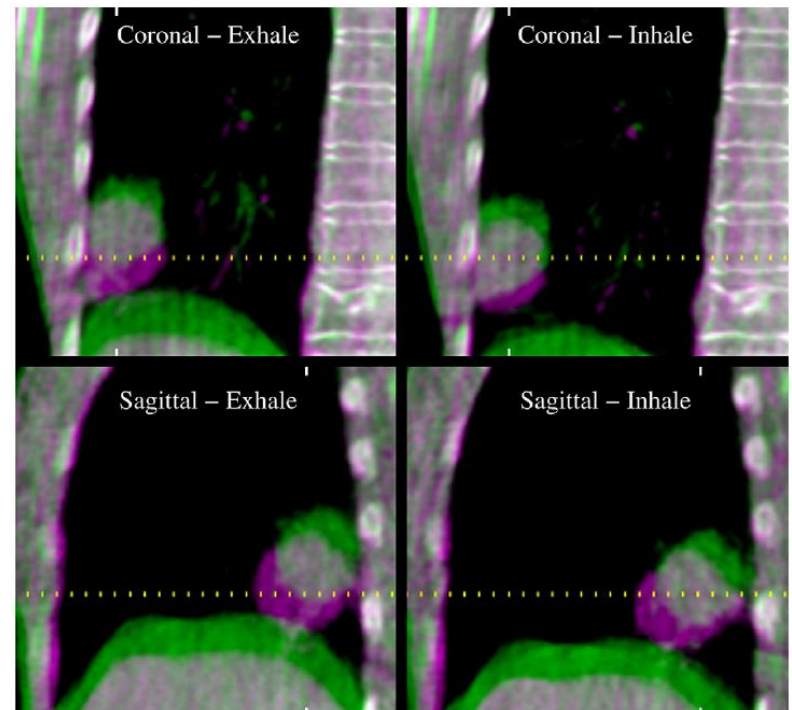
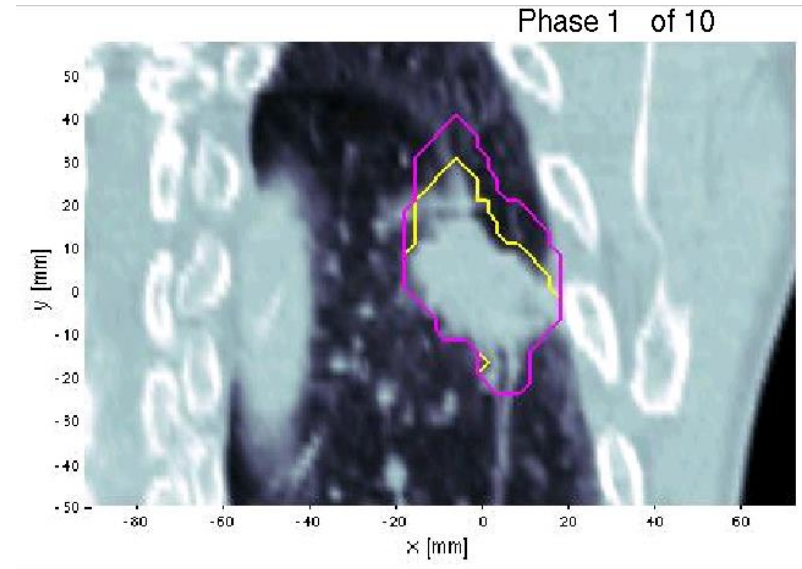


Rasch et al. Radiation Oncology 2010, 5:21

R.J.H.M. Steenbakkers et al. / Radiotherapy and Oncology 77 (2005) 182-190

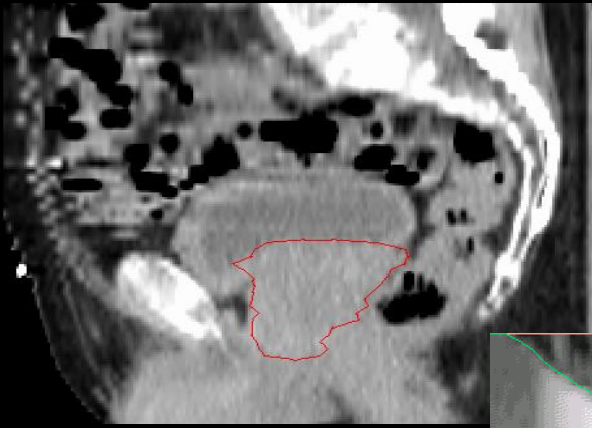
4D – Intrafraction motion (lung)

- 4D CT
 - feasibility of 4D CT
 - 4D also at treatment machine (4D CBCT)



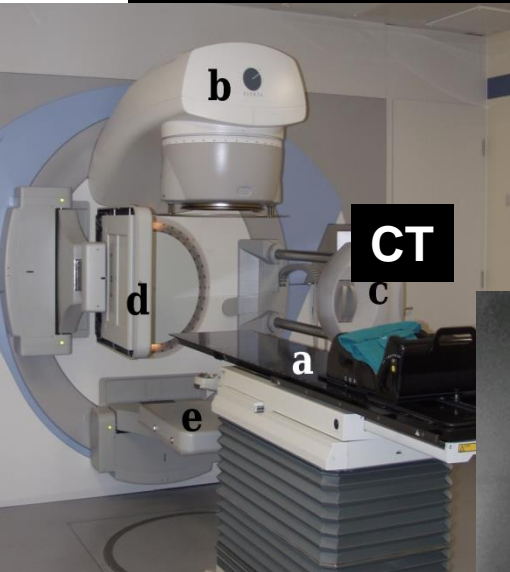
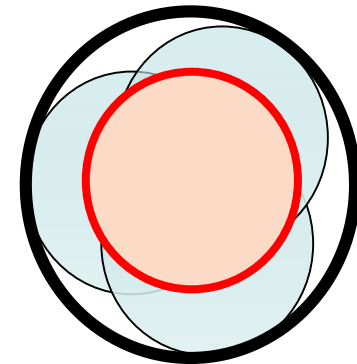
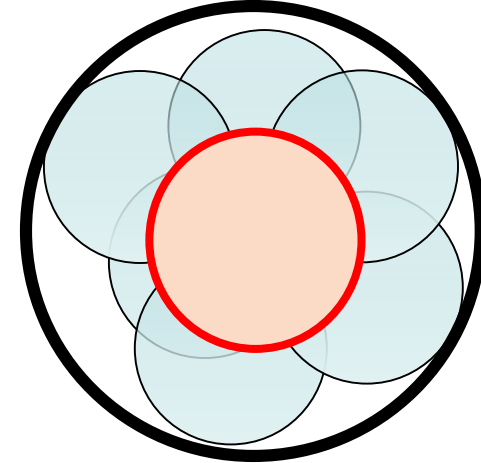
„dancing PROSTATE“
„Amsterdam prostate waltz“

Cone beam CT am Linac

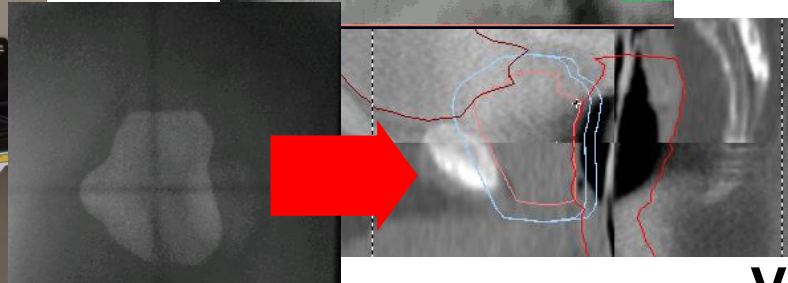
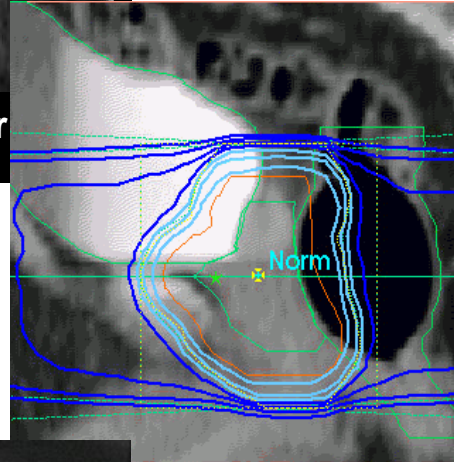


Courtesy B. Mijnheer

Intrafraction motion
reducing safety-margins
by use of IGRT (CTV-PTV/IORV)
from 5 to 2-3 mm
from 10 to 7 mm



CT

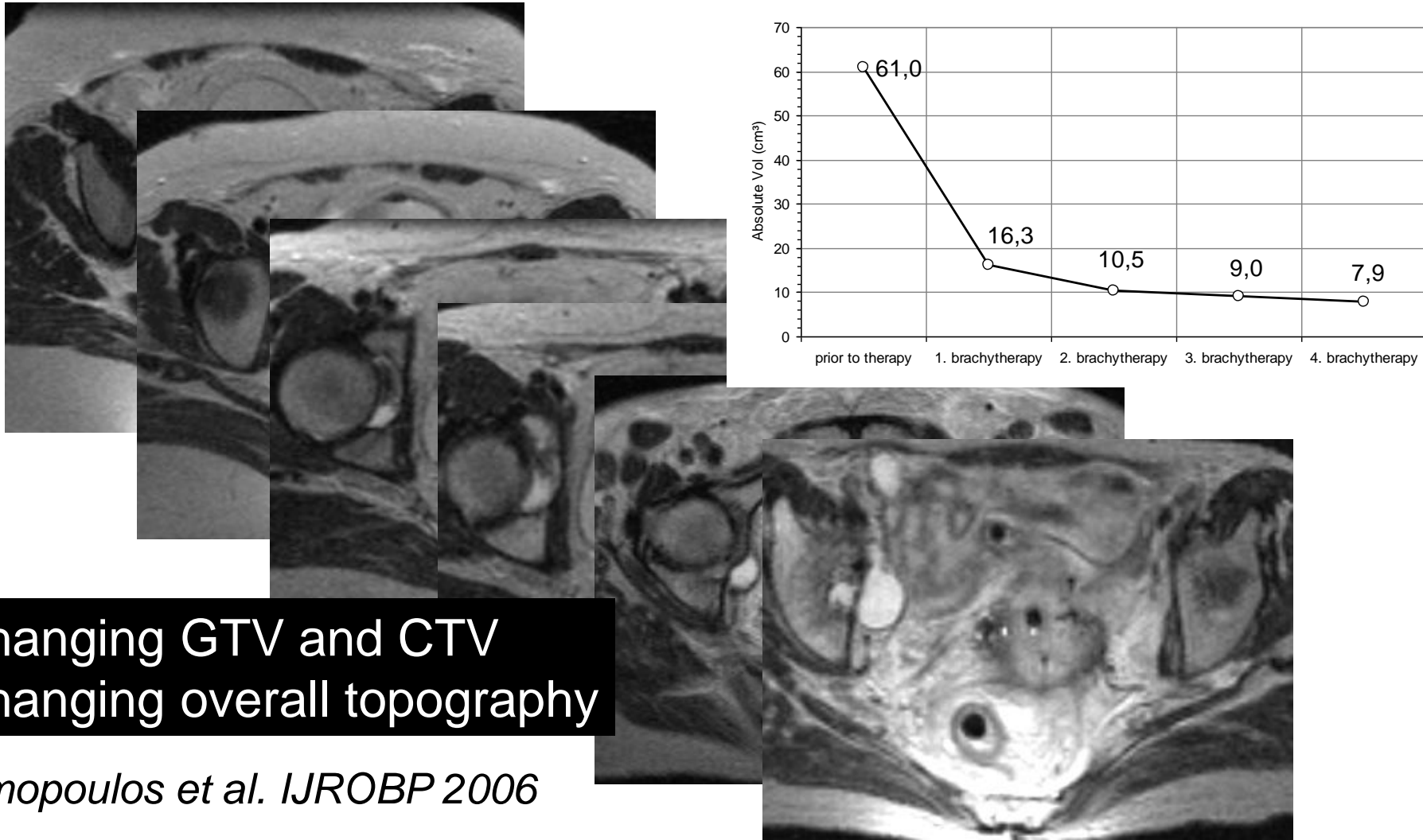


„Vienna prostate waltz“

Linac A
Image guided Radiotherapy

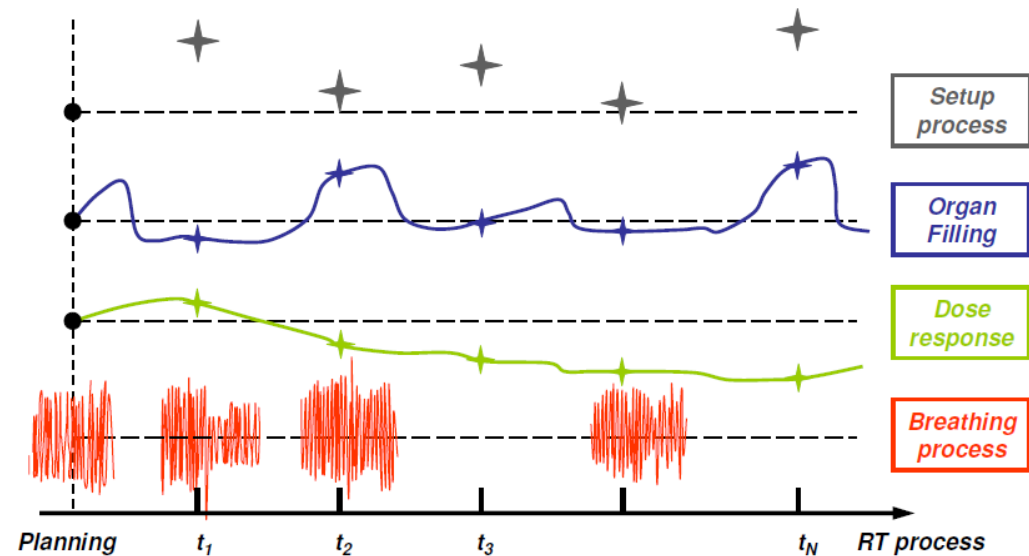
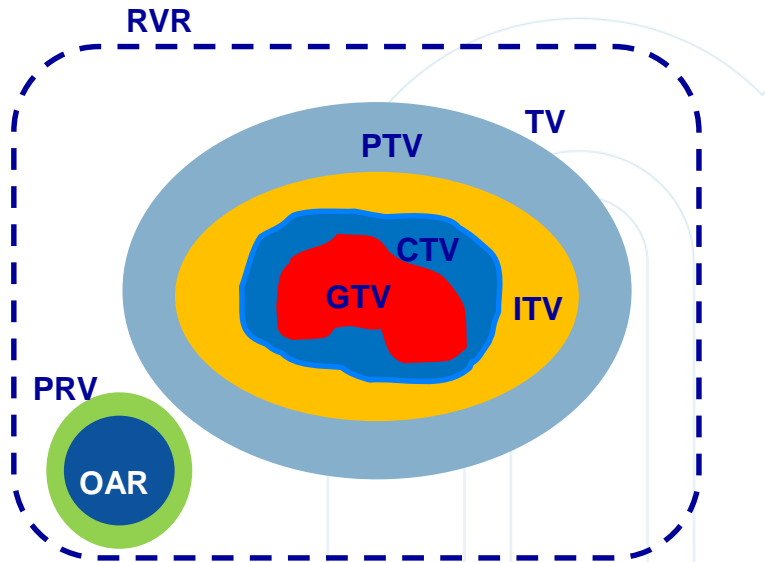
4D – Interfraction change (cervix)

- MRI: Initial tumour extension (3D RT)
- pattern of spread and response (4D RT)
- for adaptive MRI based radiotherapy (BT)



Changing GTV and CTV
Changing overall topography

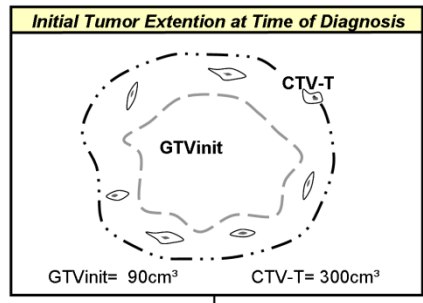
ICRU volume concepts (from 62 to 83 and 89)



- ICRU concepts have been traditionally based on morphology/anatomy
- Margins account for temporal effects
- Concepts are in transition with subvolumes for GTV defined based on functional imaging, and/or GTV response. . .

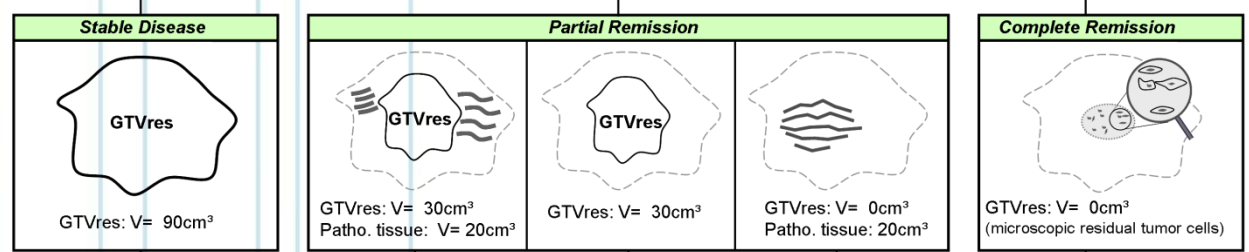
Target concepts imply structure boundaries !

change in tumour volume and tumour configuration during treatment

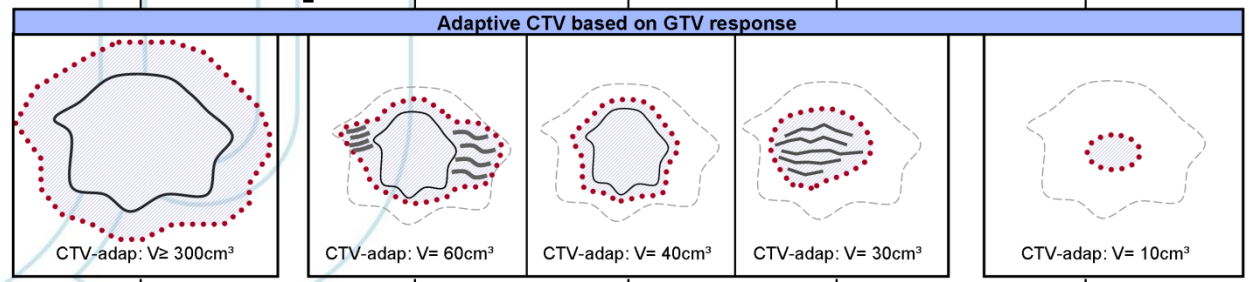


TREATMENT

Patterns of GTV Response



Corresponding various patterns of



ADAPTIVE TREATMENT

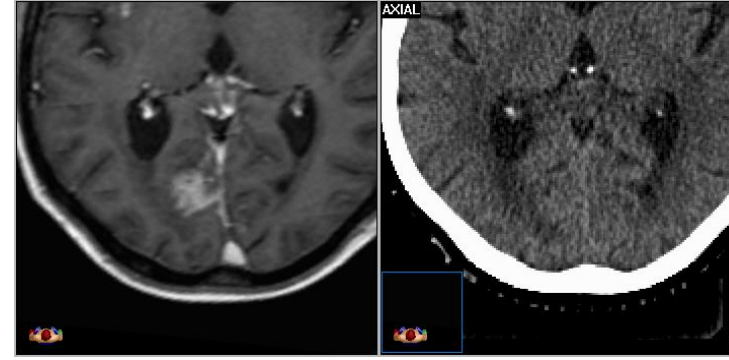


Various patterns GTV response

Corresponding various patterns of adaptive CTVs

ICRU/GEC ESTRO
Report 89
Fig 5.3

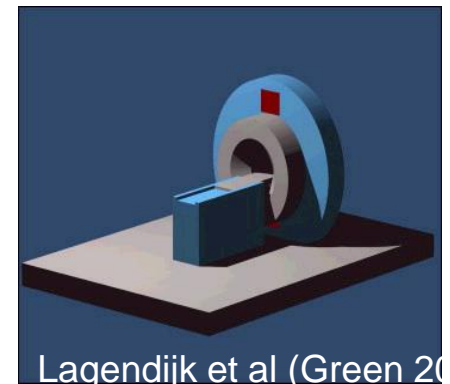
Limitations of CT



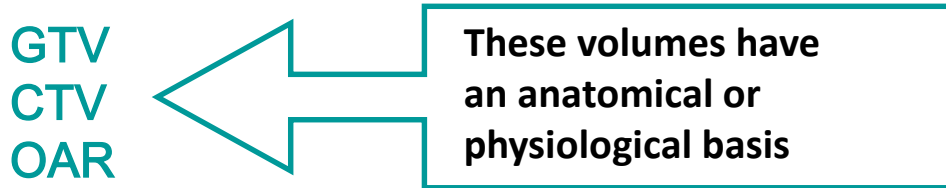
- CT in many anatomical sites:
large inter- & intra-observer variability
- MRI provides improved
soft tissue contrast with better visibility
and a large amount of sequences

Some remarks on MRI

- image distortion evaluation (MRI)
- image co-registration necessary (?)
- MRI+Linac



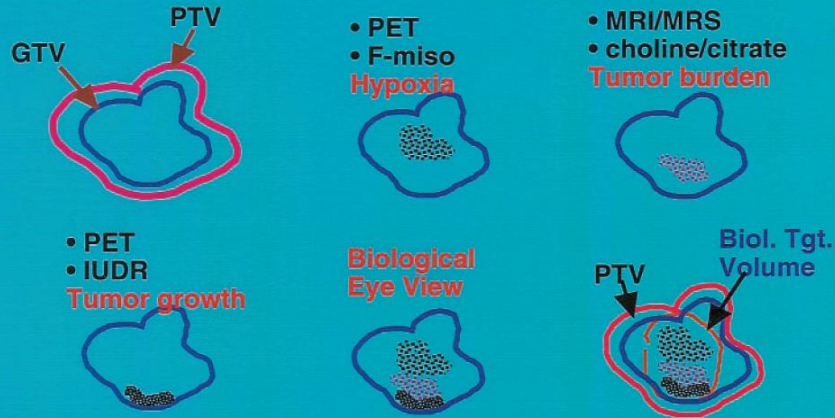
Volume definition



- Selection and delineation of the CTV and the OAR is a medical decision,
 - results from a clinical judgment involving many factors, e.g pathology and imaging findings, imaging.
- Delineation of the GTV and the CTV should be independent of the irradiation techniques, and influenced only by oncological considerations

(D3) Determine biological attributes „bio-imaging“- multimodal imaging

Biological Target Volume?



Ling et al. 2000

Homogenous dose distribution
assuming random distribution
of cancer stem cells over tumour

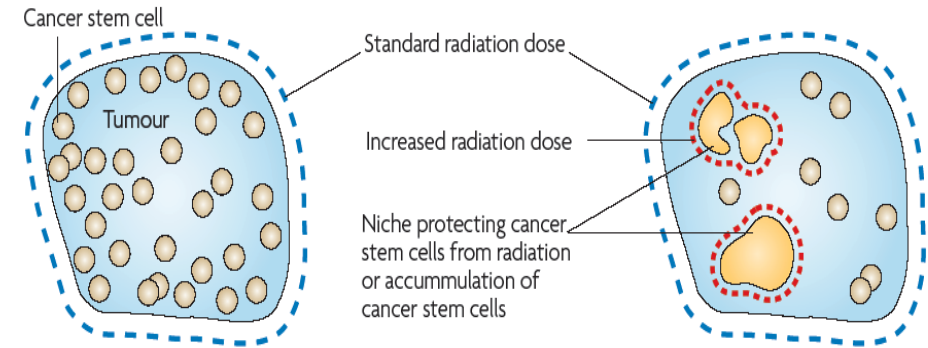
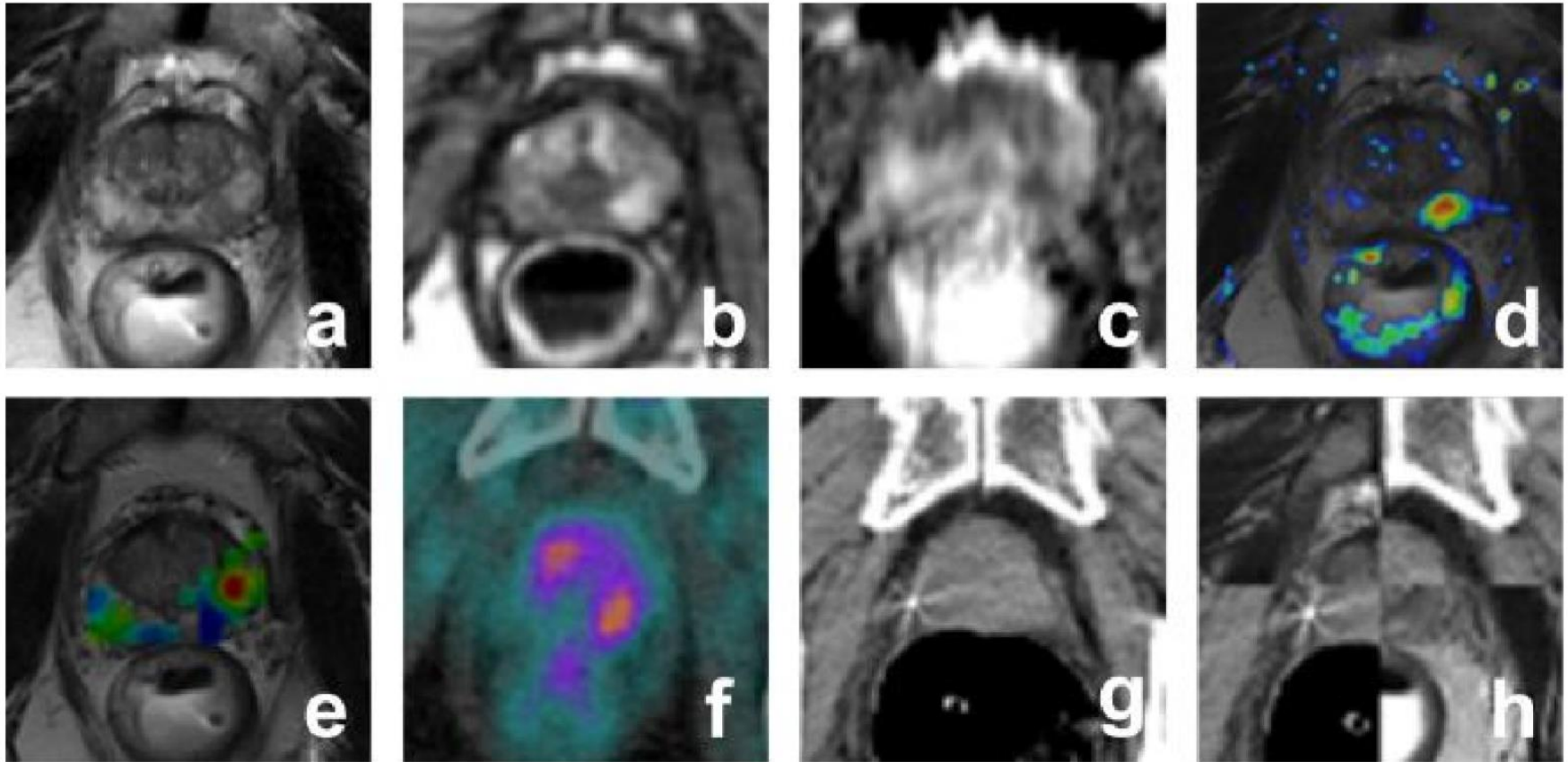


Figure 3 | The potential importance of stem cell niches for radiotherapy treatment planning.
a | Standard treatment plans would deliver the irradiation dose with a safety margin homogeneously

Baumann et al. 2008

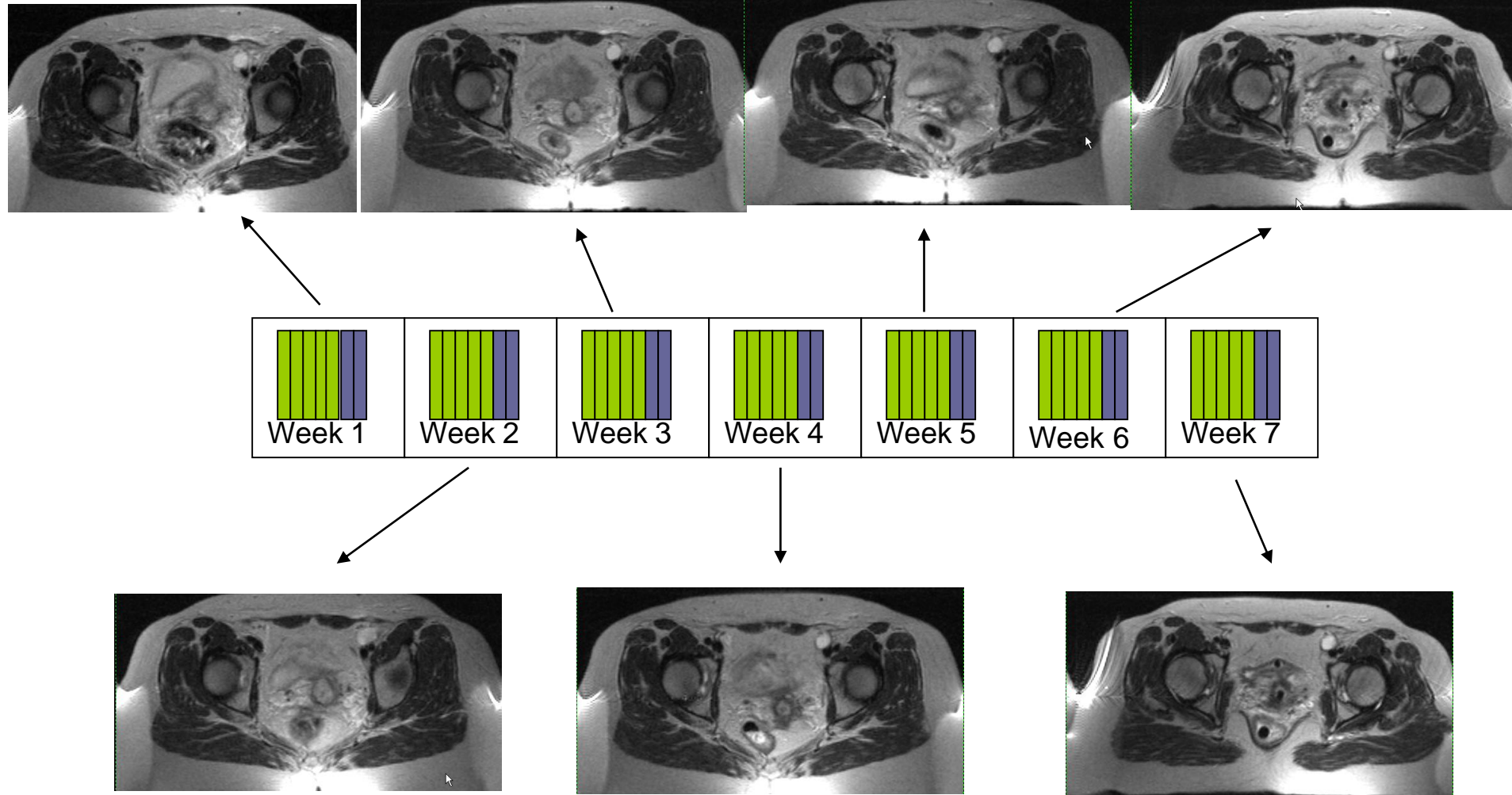
- Tumor hypoxia (H&N, Cervix...)
- Angiogenesis – microvessel density and perfusion (MRI)
-Varia.....Lactate etc.....

mp-MRI.



Supplementary Figure S1: Example of mp-MRI and PET findings for one patient: T2w (a), DCE T1w (b), ADC map (c), K_{trans} map (d) and MRS (Cho+Ci)/Cr ratio map (e). The lesion is visible in the peripheral zone at the right posterior part of the gland. For comparison $[^{11}\text{C}]$ Acetate PET-CT (f), planning CT (g) and planning CT fused with T2w MR image (h) are shown as well. [1]

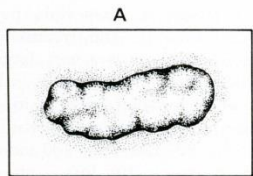
Determine „biological“ attributes from response...



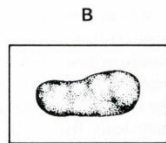
...with repetitive morphologic imaging

(D4) Dose distribution design

- 2D radiotherapy
- 3D conformal radiotherapy
- Adaptive Radiotherapy („shrinking field technique“)
- IMRT
- VMAT/Tomotherapy
- Dose Boosting/Painting (non uniform)

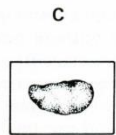


Widely around gross tumor.



(at 5 weeks, 5,000 rads)

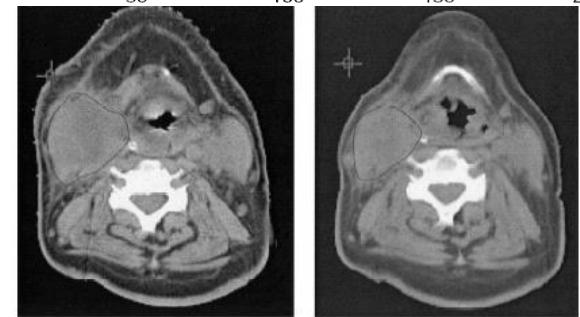
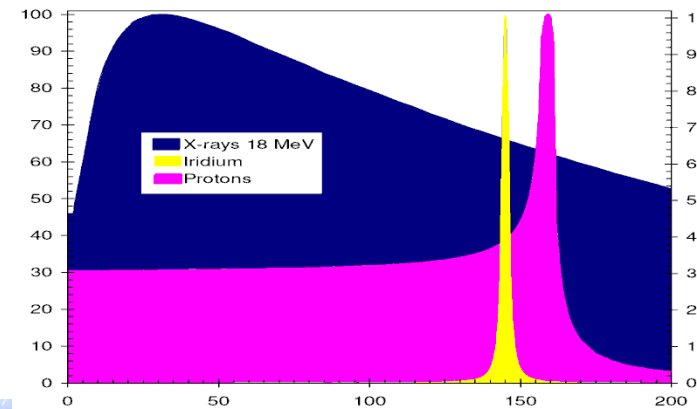
Covers gross tumor with some margin. Usually will carry to 7,000 rads total dose.



(at 6 or 7 weeks)

For very infiltrating tumor or large nodes to be treated by external irradiation only, additional 500-1,000 rads to 7,500-8,000 rads total dose.

Fletcher 1980

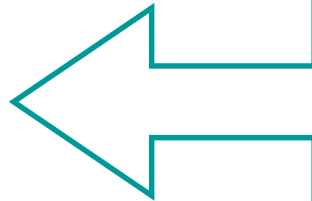


start of treatment – 3 weeks after start
Barker et al 2006

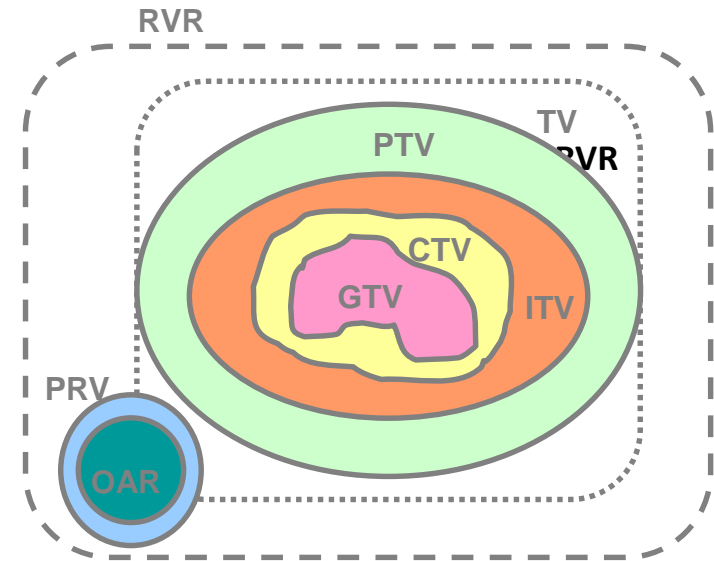
- Image guided Brachytherapy

Geometric concepts

PTV
PRV
ITV





introduced to ensure
that dose delivered to CTV &
OAR match the prescription &
constraints.



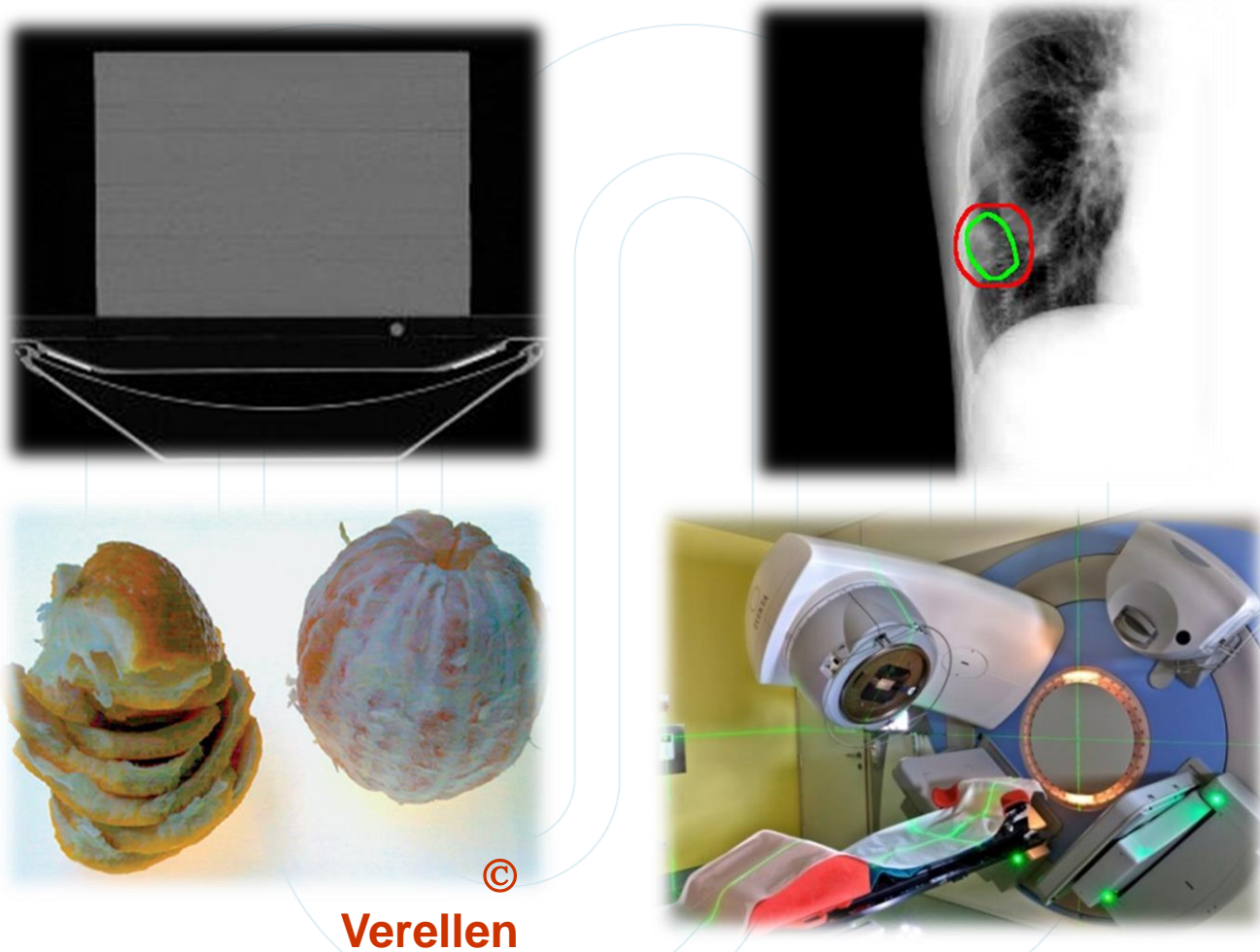
- These volumes may be constructed on TPS automatically with an appropriate margin

Margins

- Most important for clinical radiotherapy.
- Depend on
 - CTV/OAR motion  internal margins (ICRU 62)
 - patient set-up and beam alignment  external margins
- Margins may be non-uniform but should be three dimensional.
- Joint assessment of radiation oncologist and medical physicist
- A reasonable way of thinking would be:

“Choose margins so that the target is in the treated field
at least 95% of the time.”

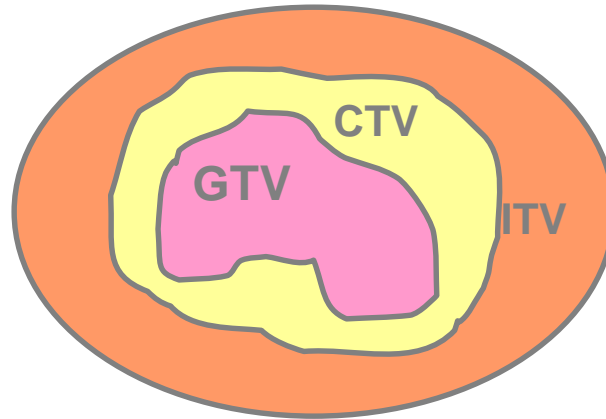
Margins in RO and Image Guidance



Geometric uncertainties are commonly accounted for by margins

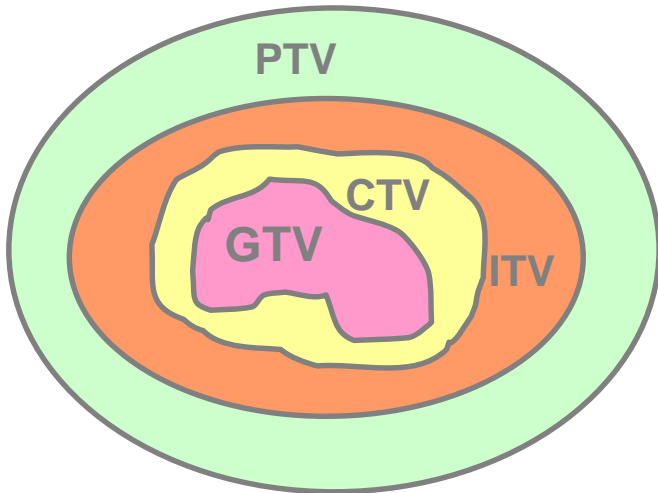
Internal Target Volume (ITV)

(complex R&D issues)



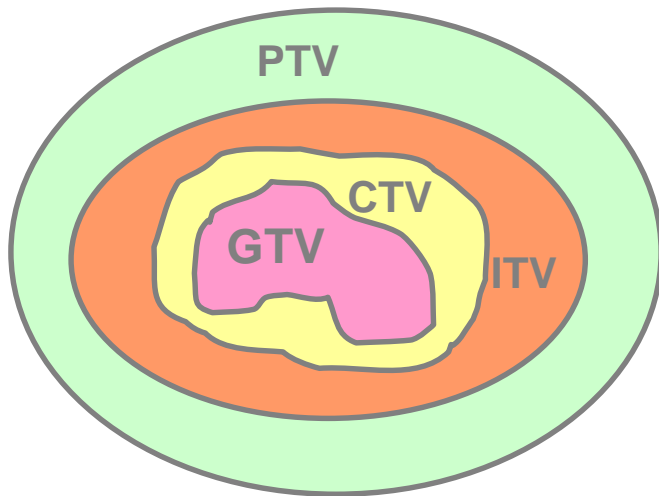
- ICRU Report 78: “In practice, it might not be necessary to explicitly delineate the ITV, but the IM (as well as the SM) must be taken into account when delineating the PTV.”
- ICRU Report 83: “The ITV is considered an optional tool in helping to delineate the PTV.”

Planning Target Volume (PTV)



- In contrast to the CTV a geometrical concept.
- It is defined to select appropriate beam arrangements, taking into consideration the net effect of all possible geometrical variations, in order to ensure that the prescribed dose is actually absorbed in the CTV.
- The PTV includes the internal target margin and an additional margin for set-up uncertainties, machine tolerances and intra-treatment variations.

How to define PTV ?



- Analyze ALL uncertainties and use appropriate margin recipe
- Systematic errors (Treatment preparation): setup error, organ motion during planning CT, delineation errors, equipment calibration errors
- Random errors (Treatment execution): inter- & intra-fraction variation

Margins in photon and proton RT

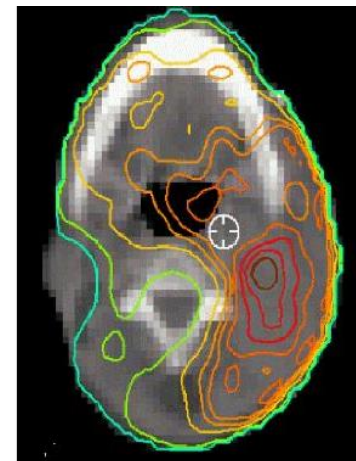
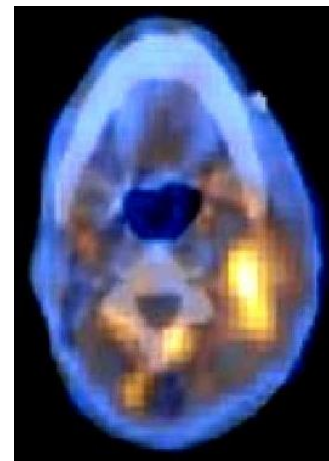
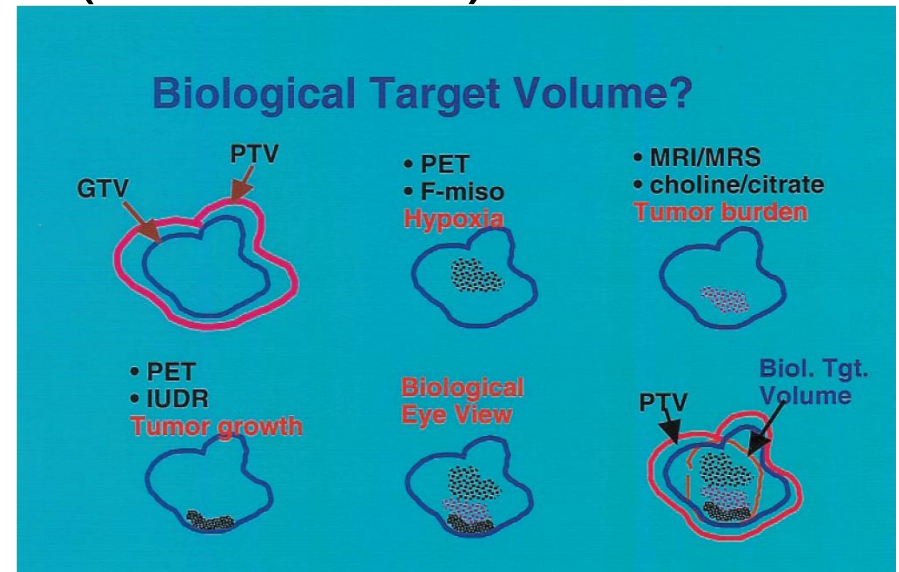
(complex R&D issues)

- **Report 83** provides margin recipes for PTV and PRV in photon therapy (ref. van Herk, Ten Haken and McKenzie)
- **Report 78** „PTV requires different margins lateral, distal and proximal to the CTV. “
- **“Daily practice” in PT:** Beams can be designed directly for the CTV, taking into account the need for internal and external margins within the aperture design, without reference to a PTV beams.
 - Nevertheless, PTVs must be defined since they are required for reporting purposes.

The delineation of the PTV is a required part of the treatment prescription

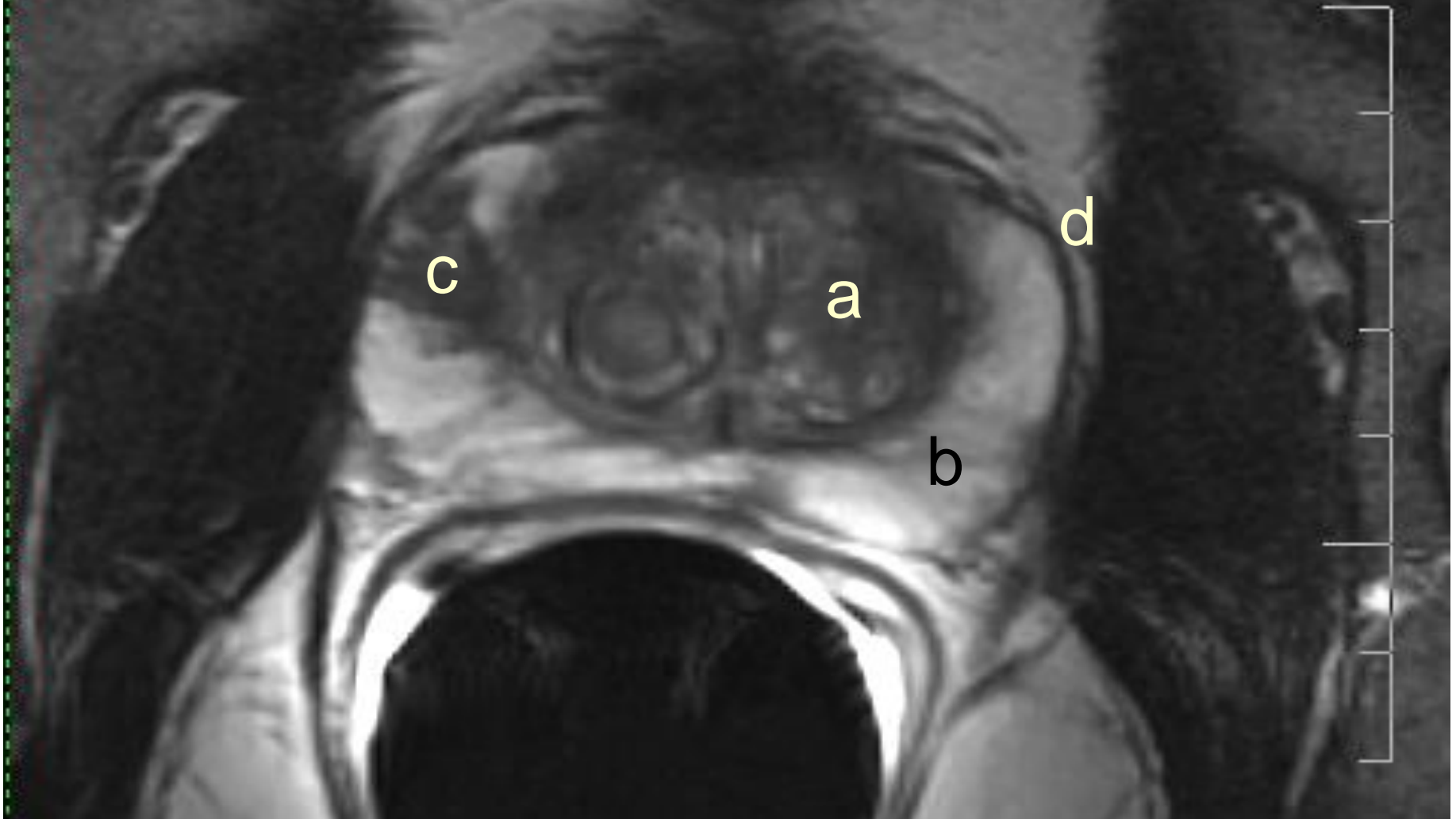
Dose Boosting and Painting

- coined by CC Ling et al (Red 2000)
- challenges dogma of homogeneity in target
- functional imaging for volumetric map of radiobiological factors
- lack of clinical trials “endpoints”



MR imaging before treatment (T2)

(a) Central lobe, (b) peripheral zone, (c) tumor, (d) prostate.



Definition of target volumes and dose prescription

CTV Low Risk

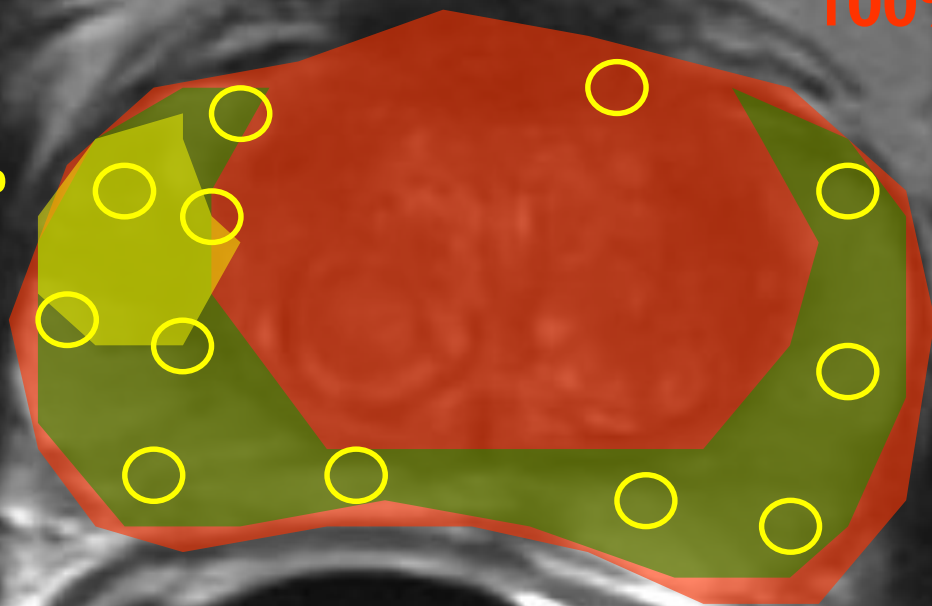
CTV Intermediate Risk

CTV High Risk

200%

100%

150%



Dose Painting: where?

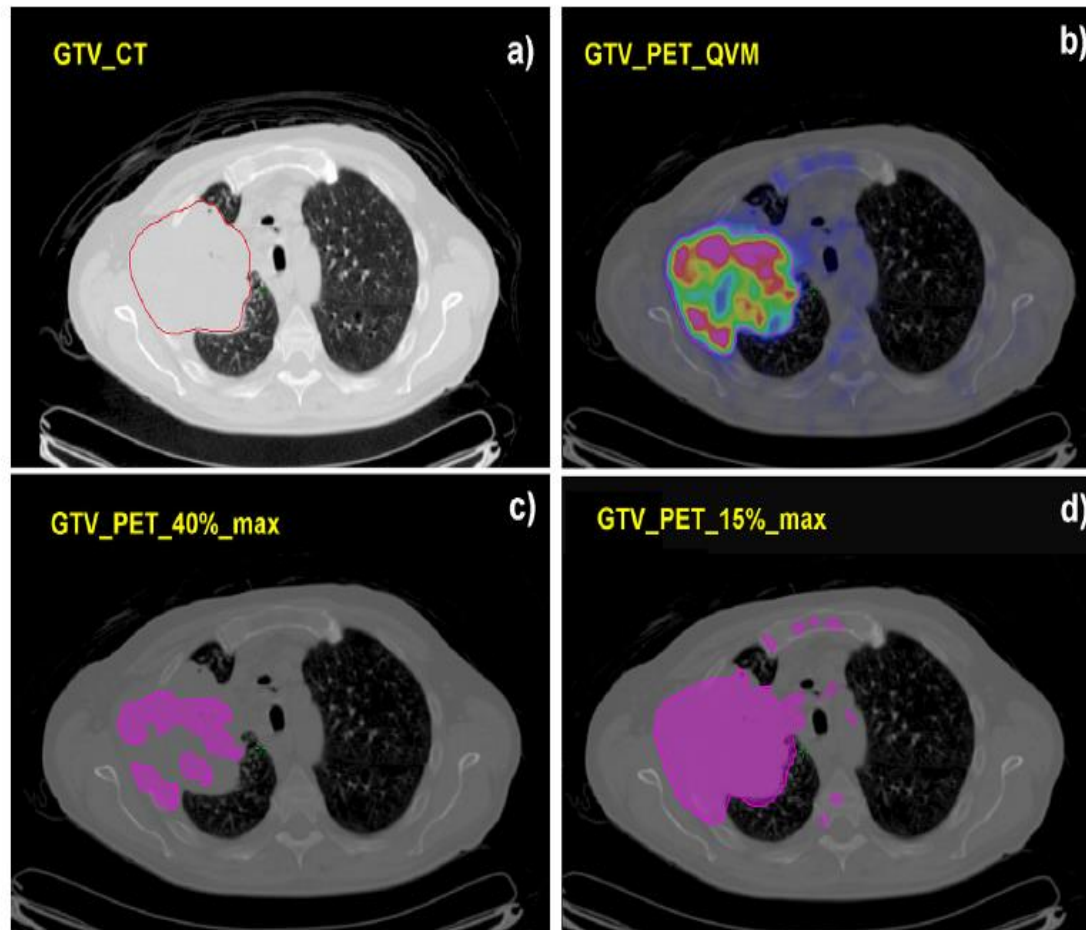
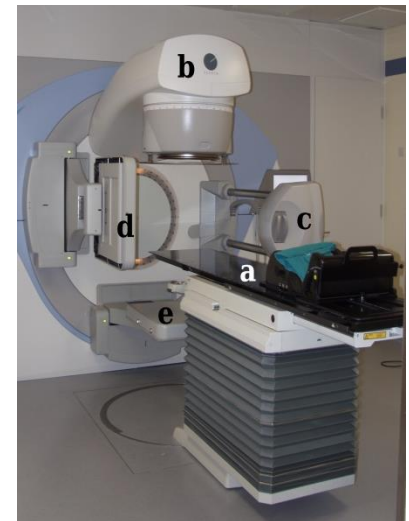


Fig. 2. Segmentation of target volumes. (a) Computed tomography-based gross tumor volume (GTV_CT) outlined by radiation oncologist. (b) Positron emission tomography-based gross tumor volume (GTV_PET) outlined by same radiation oncologist. QVM = qualitative visual method. (c) Positron emission tomography-based volume, outlined using gross tumor volume 40% maximal uptake method (single maximum) (GTV_PET_40%_max). (d) Positron emission tomography-based volume, outlined using gross tumor volume 15% maximal uptake method (single maximum) (GTV_PET_15%_max).

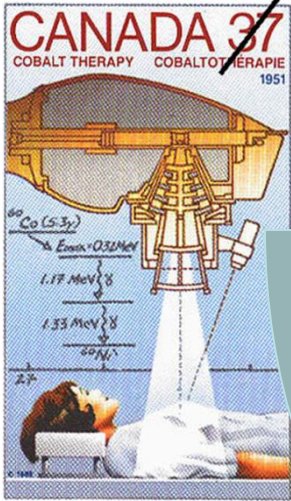
(D5) Dose delivery assurance („IGRT“)

- near real-time imaging during treatment delivery
- Ideally:
 - 3D volumetric study of soft tissue structures
 - efficient acquisition and comparison
 - process for clinically meaningful intervention
- commercial systems only partly achieve these needs



(Short) History of IGRT

1951: The very beginning ...
intended +/- 1cm



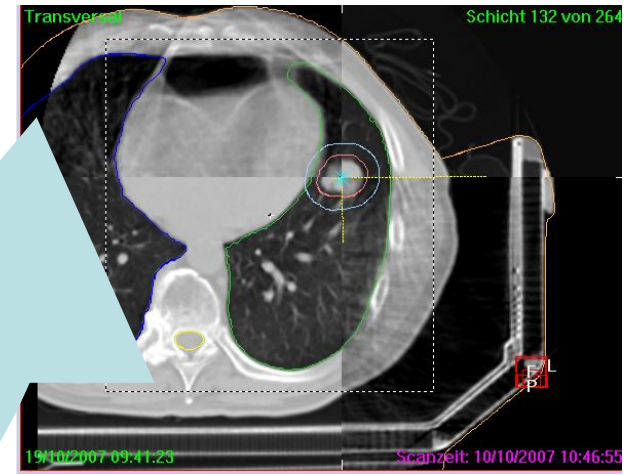
A (short) history of image-guided radiotherapy

Dirk Verellen*, Mark De Ridder, Guy Storme
UZ Brussel, Oncologisch Centrum, Radiotherapie, Belgium

Radiotherapy and Oncology 86 (2008) 4–13

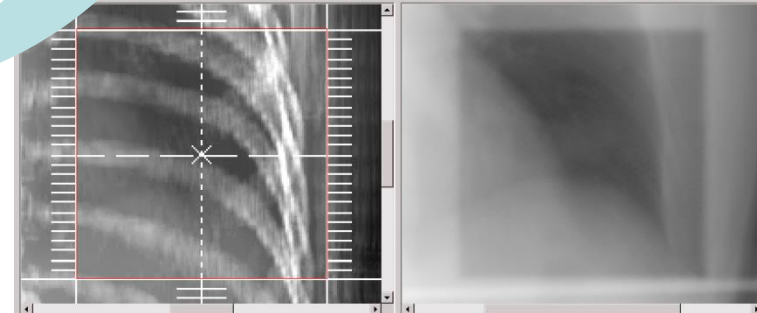
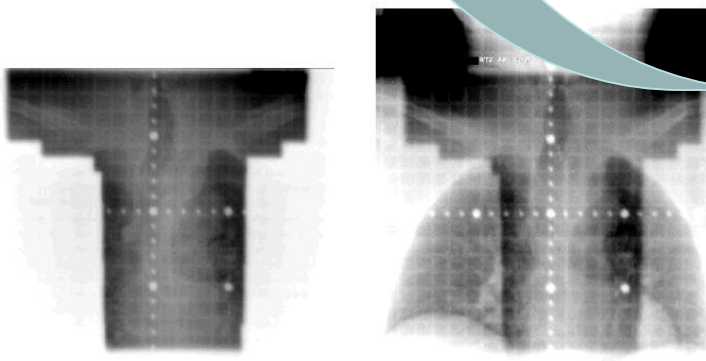
„State of the Art“
in IGRT: CBCT

achieved +/- 1mm



Since **1969**: X-ray localisation film

Since **1970**: Electronic Portal Imaging Device



Where do we need what (R&D)?

- Where there is little movement, e.g. brain, (H&N)
 - bony landmarks
 - 2D MV, kV imaging with „well-thought-out“ correction protocol probably sufficient
- Where targets move relative to bony anatomy
 - 2D imaging with radio-opaque markers
 - no info on normal tissue or tumor conformation
- Where targets move relative to OAR:
3D imaging for comparison (e.g. SRT lung, liver)
 - online imaging → systematic and random errors corrected

„dancing PROSTATE“
„Amsterdam prostate waltz“

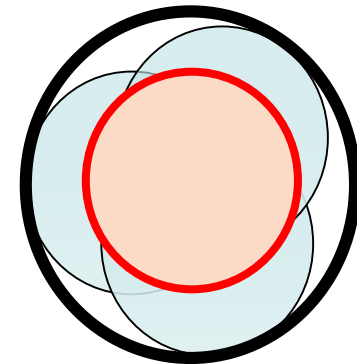
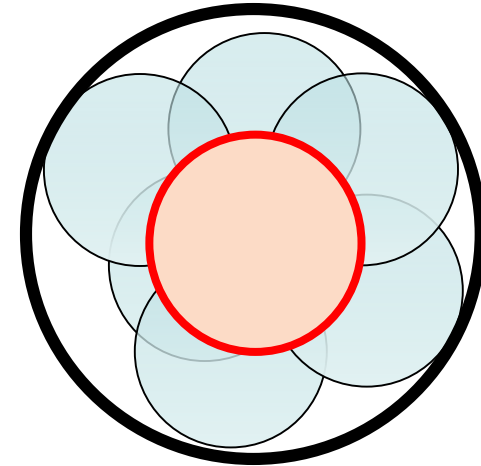
Cone beam CT am Linac



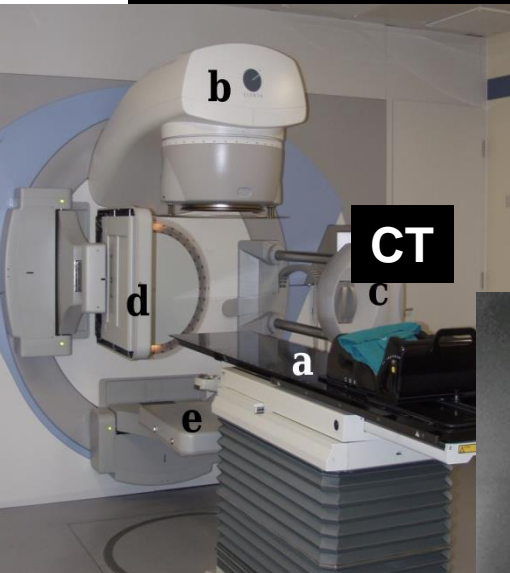
Courtesy B. Mijnheer

Reducing safety-margins
by use of IGRT (CTV-PTV/IORV)

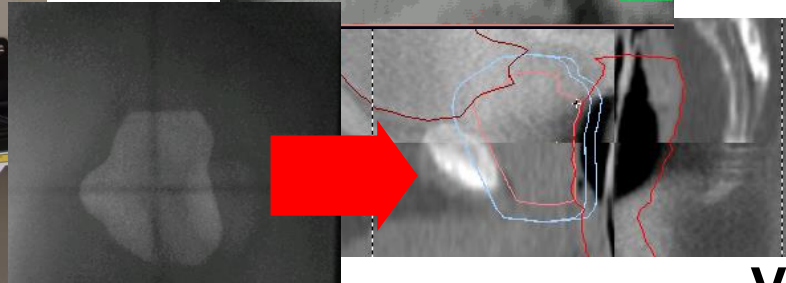
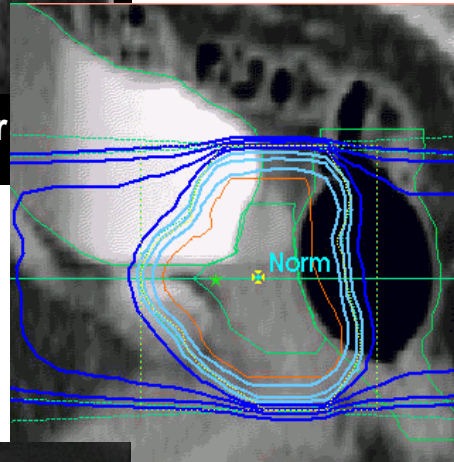
from 5 to 2-3 mm
from 10 to 7 mm



„Vienna prostate waltz“



CT



Linac A
Image guided Radiotherapy

Today's Technology for Image Guidance

- Beam quality

- MV (3 – 6 MV)
- kV (80 – 130 kV)

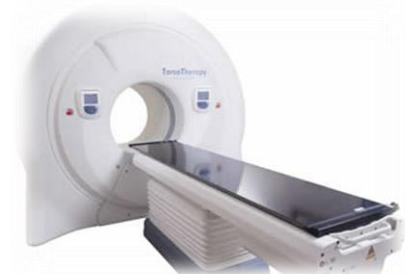
- Beam collimation

- CBCT
- FBCT

- Dimensions

- 2D
- 3D

- Rail-track-, ceiling/floor-, gantry-mounted



Current IGRT technology on/in the linac is X-ray based

Maximal conformity at maximal costs ?

Radio-Oncology

Upcoming tools and „toys“
for advanced treatment

Cyberknife

Proton
IMPT

Carbon
ions

.....

Tomotherapy

IMAT / VMAT

IGRT + IGART

Stereotactic RT

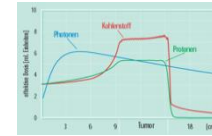
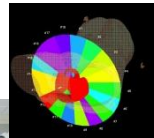
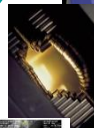
IMRT

MLC

3D

2D

Out
come



Upcoming tools and „toys“
for morphologic+biological imaging

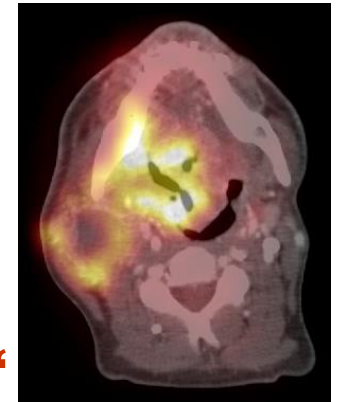


Image Guided Radiotherapy (IGRT)

- Reduction of setup and internal margins
- Reduction of side effects
- Enables dose escalation
- Room or rail-track mounted system
- Gantry or couch mounted system

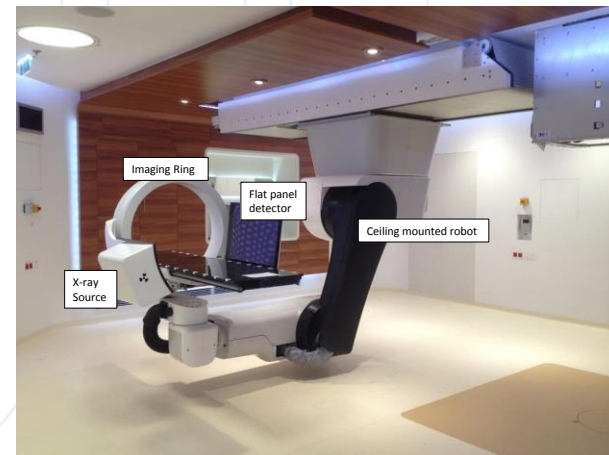
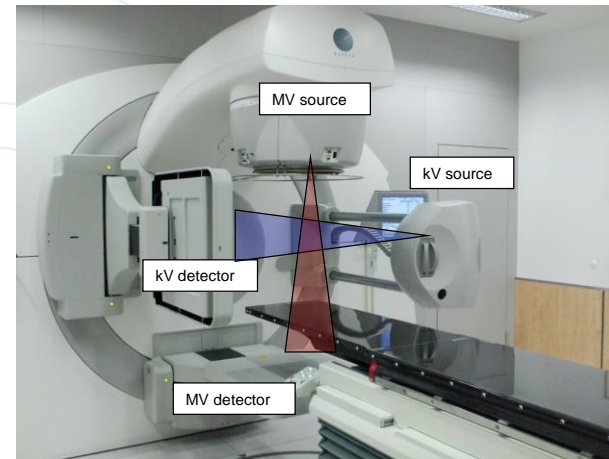
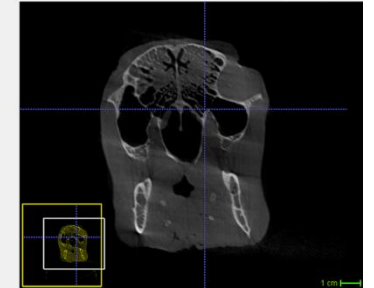
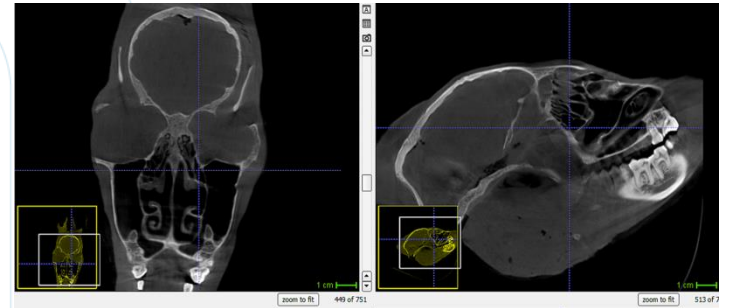


Image Guidance in proton therapy

- Example MedAustron: Imaging Ring (MedPhoton)



(D6) Deciphering treatment response

(CR, PR, SD, PD/ time pattern)

Standard:

- **Morphological changes;**
**clinical exam., light imaging (endoscopy),
US/CT/MR**

Upcoming:

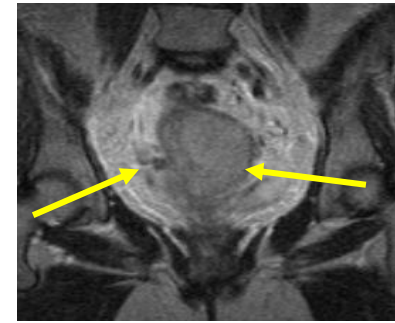
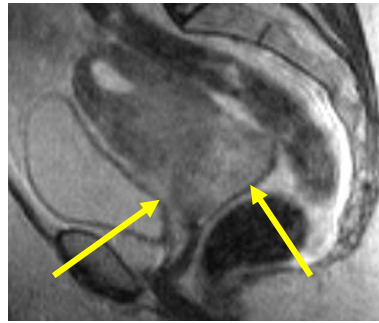
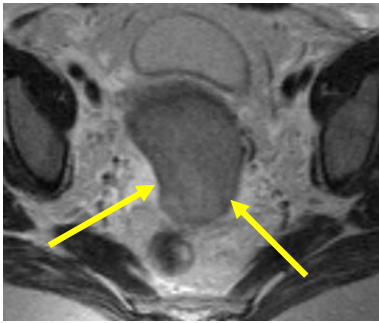
- Comparison pre- at and post treatment
functional imaging (FDG)-PET CT, fMRI

Present Developments

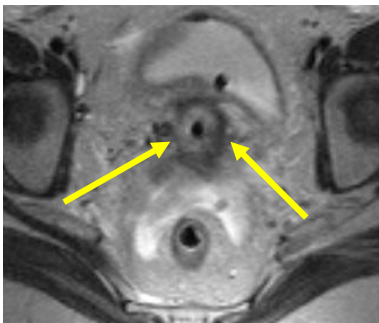
- Measurement of RT induced metabolic changes (cellular proliferation - FLT, Apoptosis,...)
- ADC (apparent diffusion coefficient) DWI,, K trans, (DCE) etc.

STAGE IIB: 5 cm WIDE, SUFFICIENT RESPONSE (12/2002)

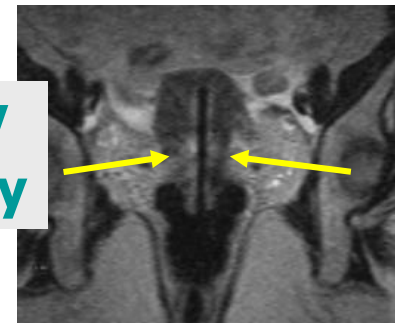
At diagnosis: 5 cm wide, 5 cm thick, 7cm high: 88 cm³



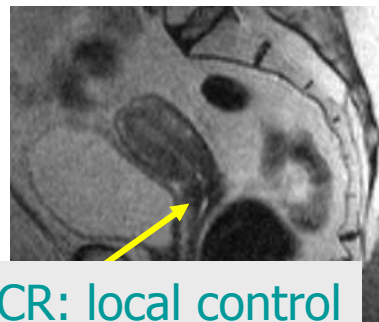
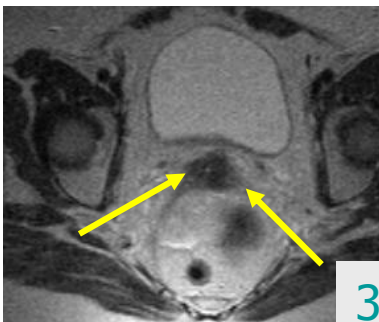
At first BT: 2 cm x 3 cm x 3 cm, 9 cm³, good remission, sufficient for intracavitary BT



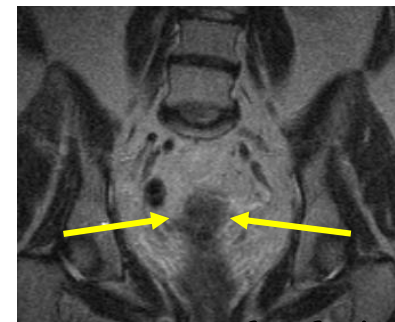
**HR-CTV
D₉₀:88Gy**



6 months after treatment: Continuous Complete Remission



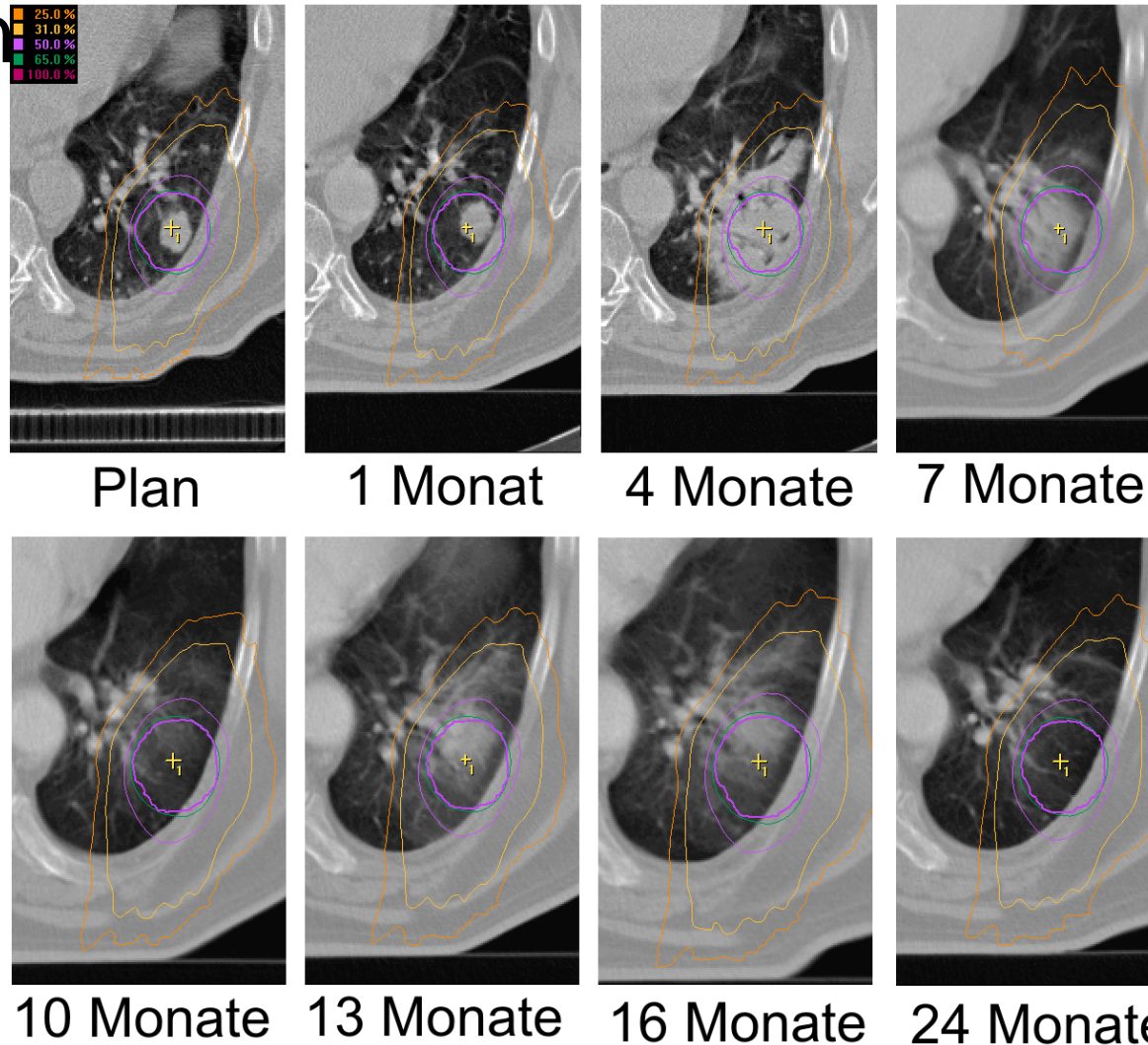
3/2005: CCR: local control



(D7) Diagnosis of outcome (after treatment)

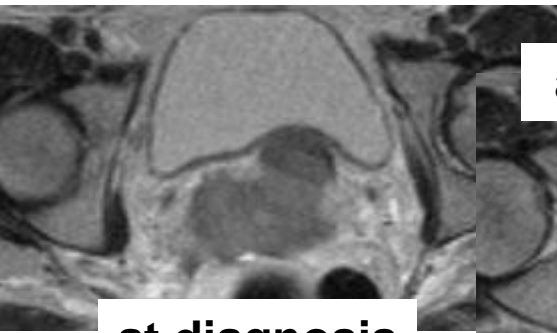
Image assessment after certain time intervals

- Tumour remission
(complete)
- Lung fibrosis
(transient)



after
Definitive
Stereotactic RT
in T1 lung cancer

**Stage IIIB, 6 cm, insufficient response (9/2000)
no adaptation of application technique
Intracavitary BT alone**



at diagnosis



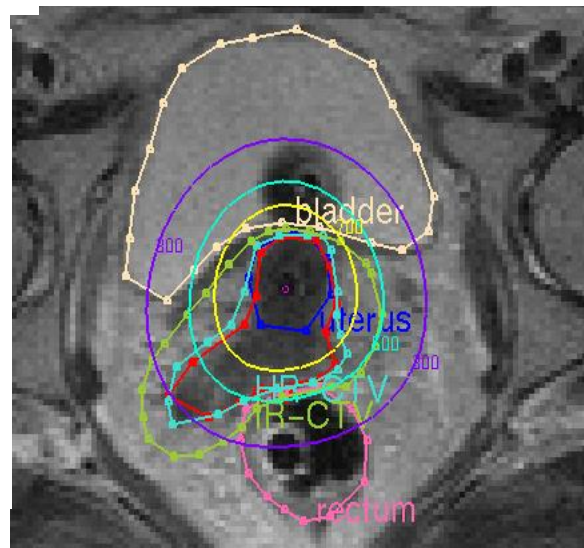
after EBRT, first BT



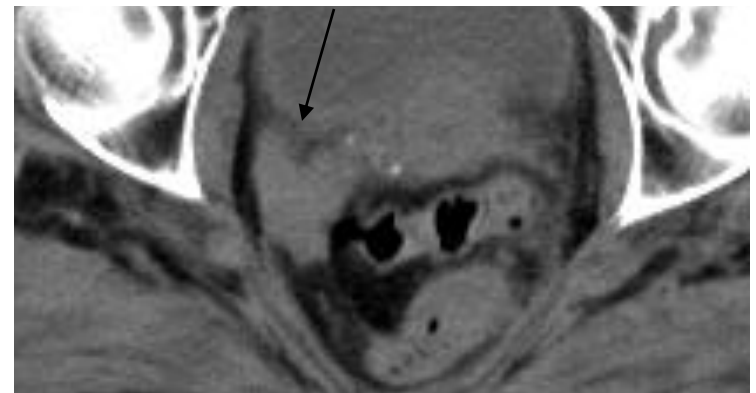
second BT



third BT



**Parametrial/pelvic wall recurrence
8/2001**

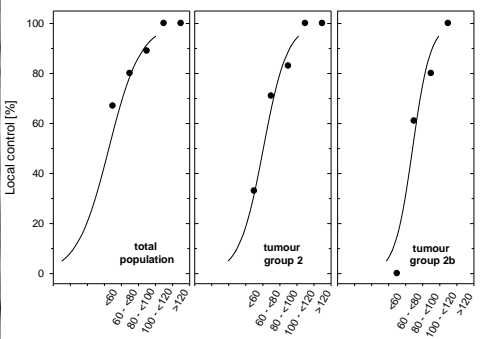
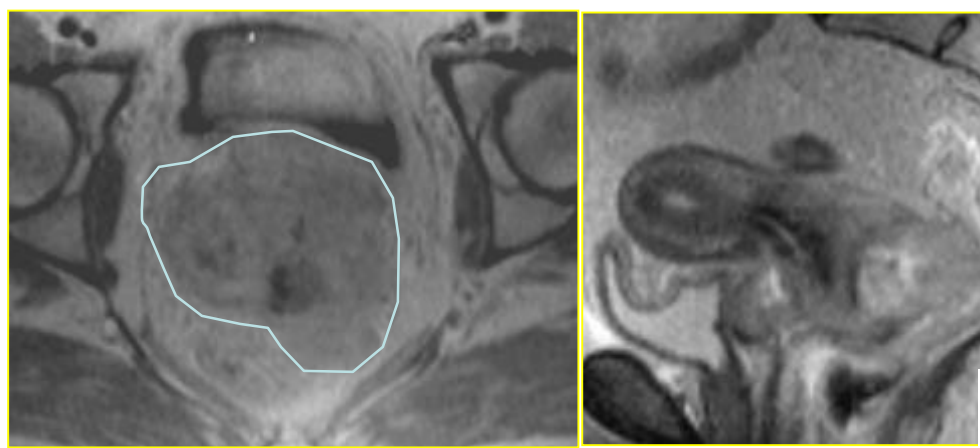


**Optimisation
based on
intracavitary BT
alone
HR CTV
D90: 67 Gy**

4D analysis of tumour spread, target coverage and recurrence

Insufficient Tumour remission (cervix cancer stage IIB): no adaptation of application technique topographical correlation to DVH parameters

Diagnosis

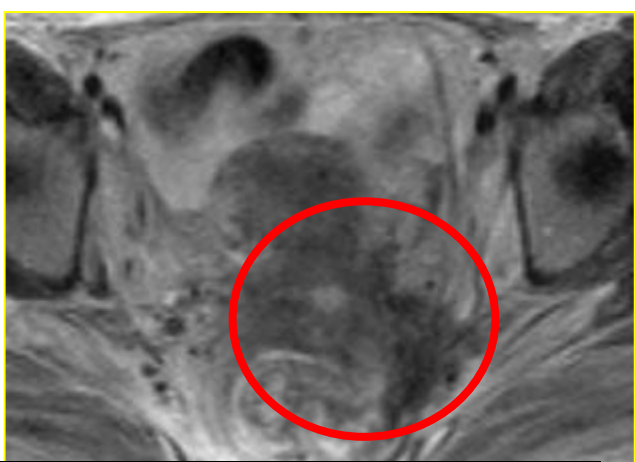
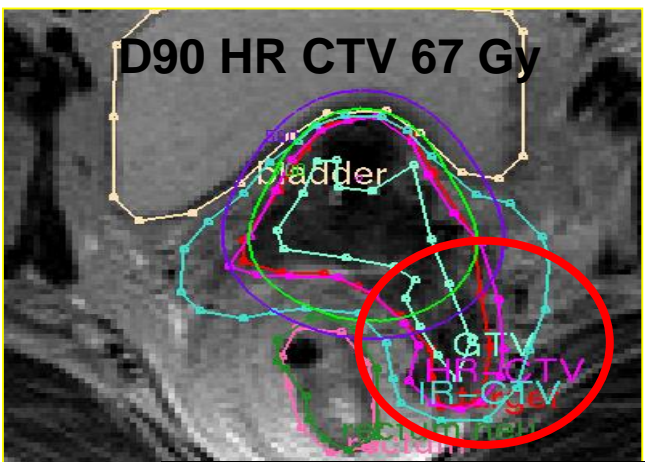
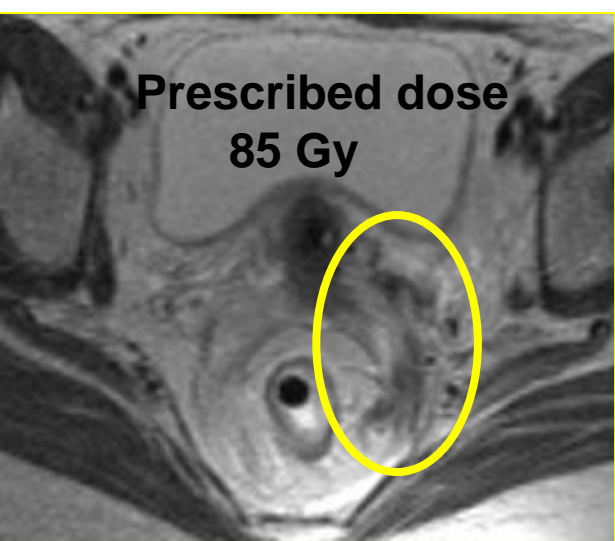


Dimopoulos et al. 2010

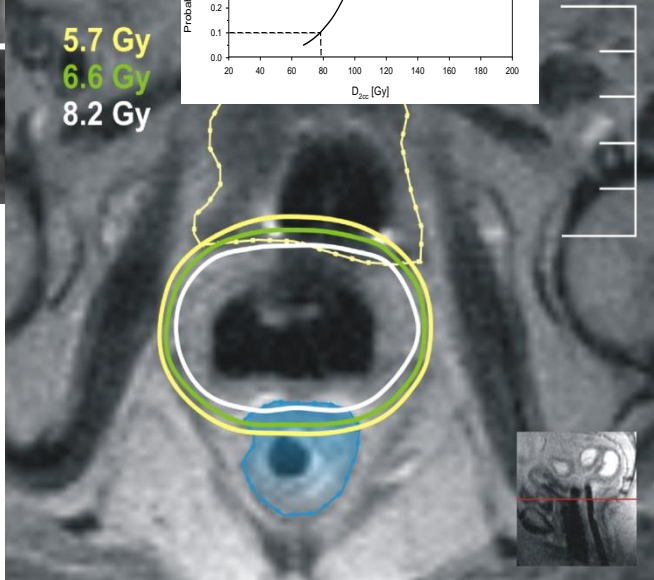
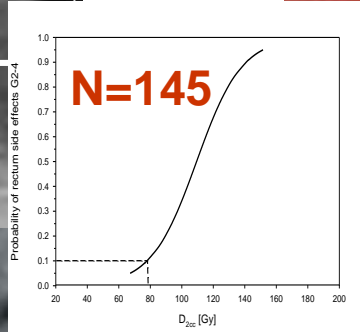
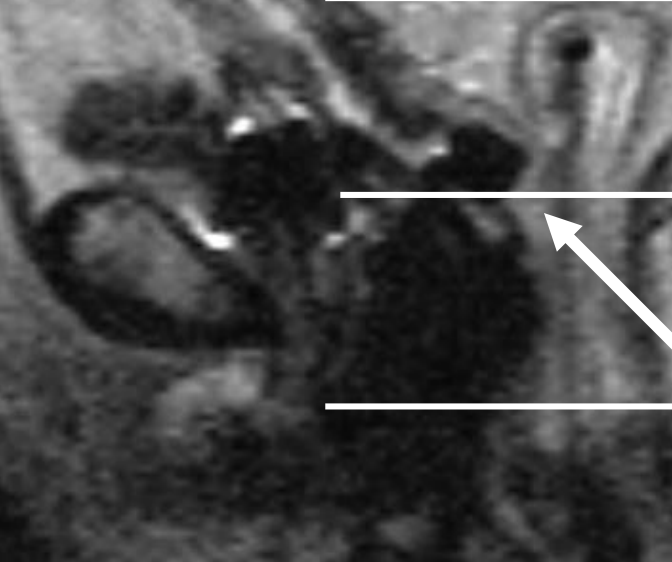
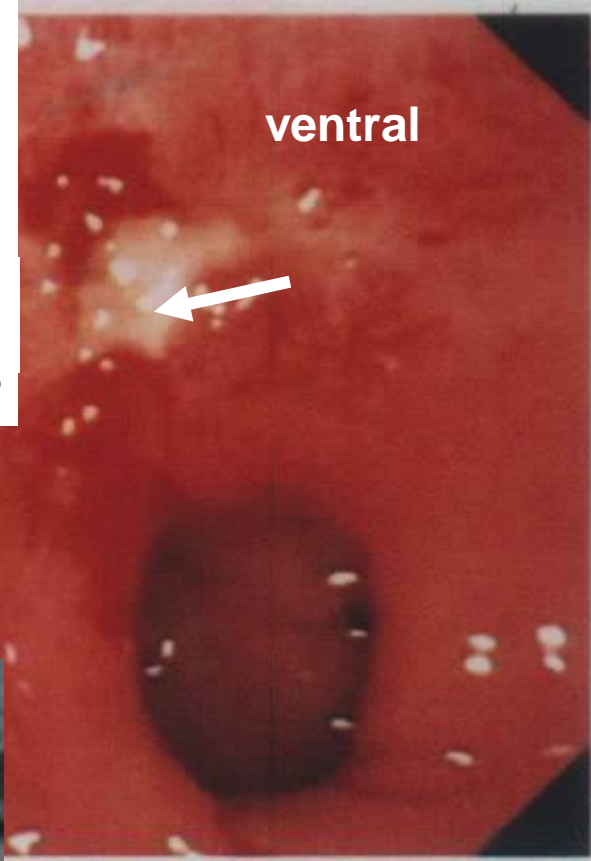
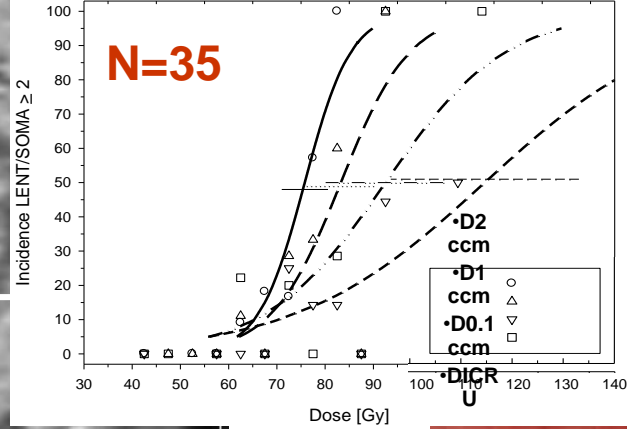
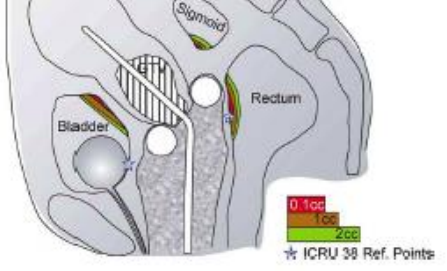
At Brachytherapy

Brachytherapy (standard)

9 mths later: Recurrence



Dimopoulos et al. GEC-ESTRO 2005

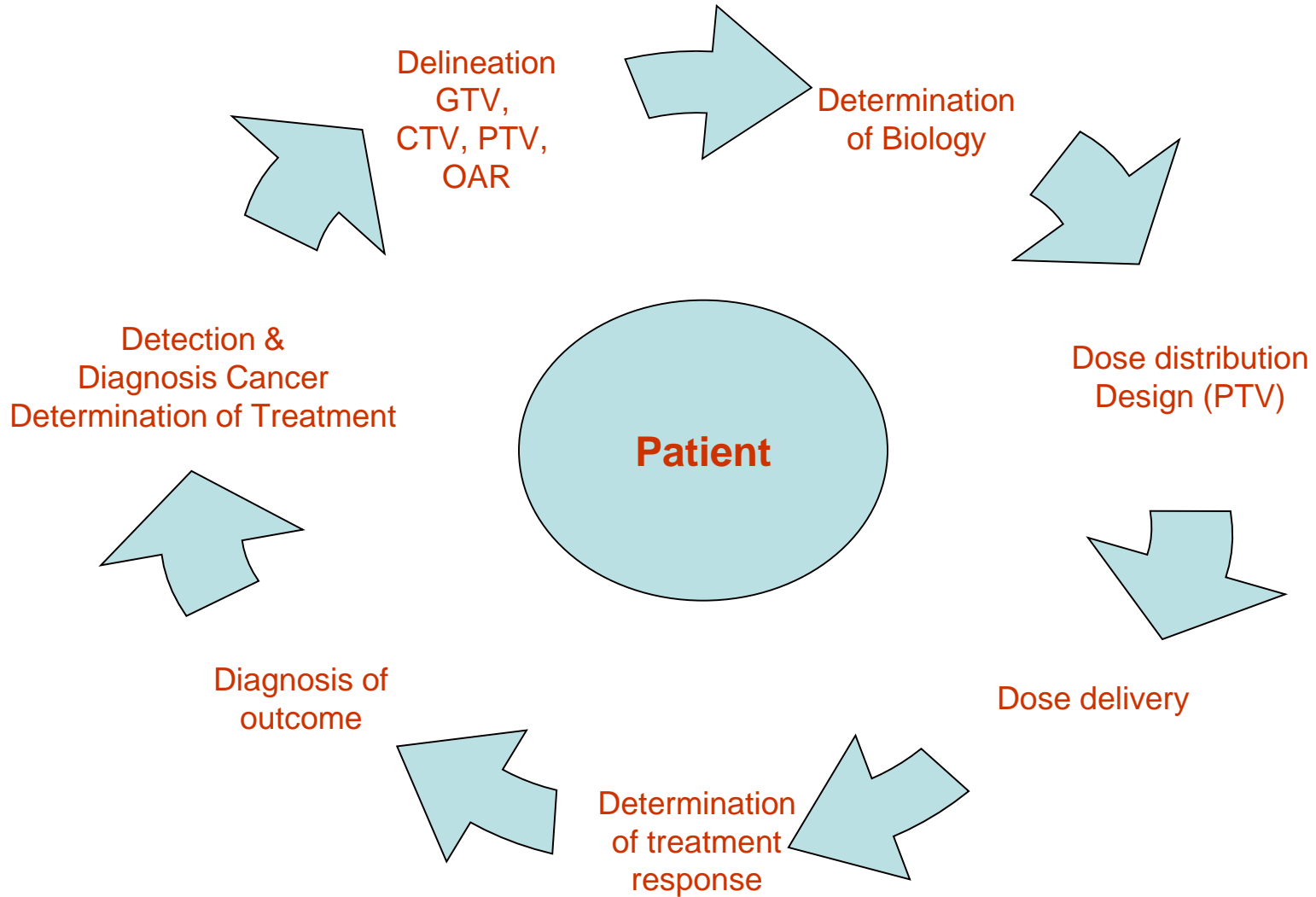


D2cc= 81 GyEQD2
D1cc= 90 Gy EQD2
D0.1cc= 108 Gy EQD2

←
high dose area
corresponding to 0.1cc

Example of one patient with rectal ulceration in distal rectum at the anterior rectal wall. The location of this ulceration corresponded to the small area of 0.1cc of rectum receiving a dose of 108 Gy EQD2.

Imaging-Cycle



IGRT: a comprehensive view (7D): activities within this overall frame

- Detection/Diagnosis cancer, Determinat. treatment
- Delineation of target (CTV/PTV), Organs at Risk
- Determining biological attributes
- Dose distribution design (PTV)
- Dose delivery assurance
- Determining treatment response
- Diagnosis of outcome (recurrence/morbidity)

Major clinical relevance

Multimodal Imaging

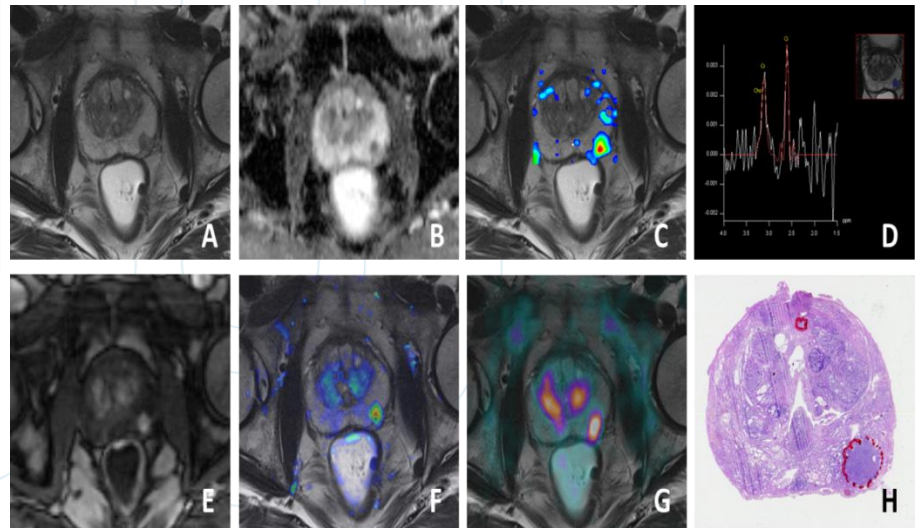
Research

Work in progress

Multiparametric imaging

- **Combining image-derived parameters to increase diagnostic accuracy**
 - improved sensitivity and specificity due to complementary information
 - reduced patients burden: cost- and time-effectives
 - suitable for diagnosis and treatment planning

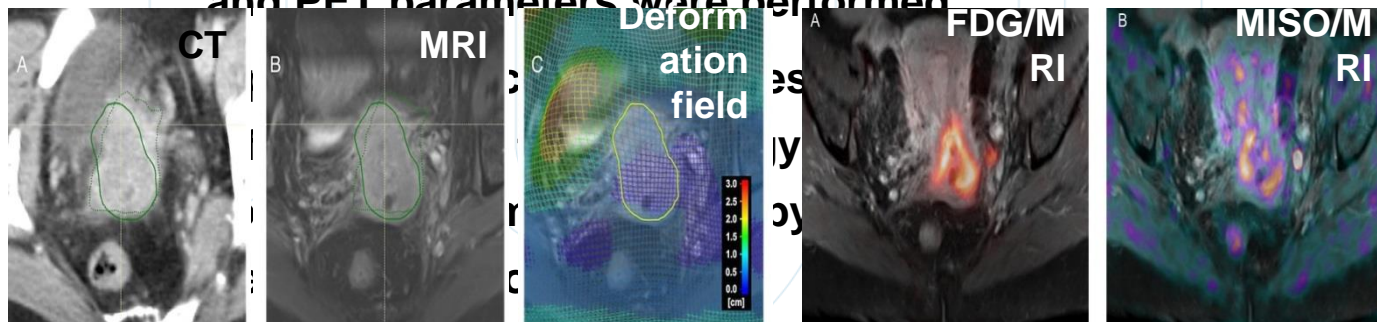
- **Multiparametric?**
 - anatomy
 - Vascularization
 - Perfusion/permeability
 - cellularity/proliferation
 - chemical composition



*Polanec S. and Andrzejewski P. et al
(submitted)*

Cervix cancer (CCa) characterization with mpMRI and $[^{18}\text{F}]\text{MISO}$ and $[^{18}\text{F}]\text{FDG}$

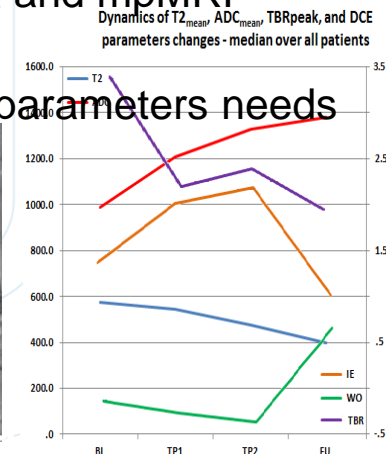
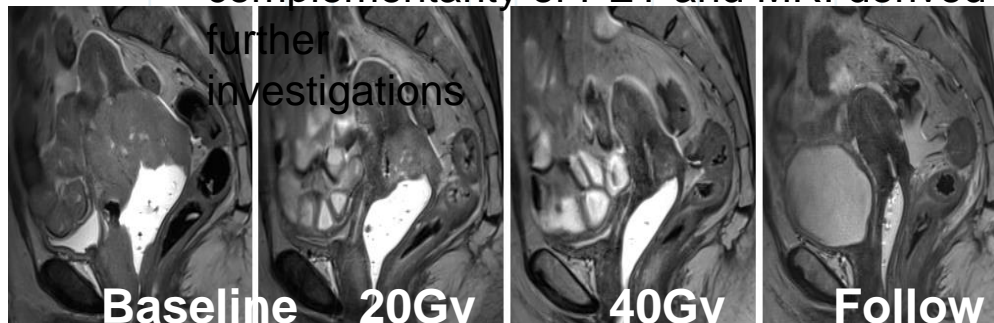
- **Dataset: 11 CCa patients scanned with mpMRI, $[^{18}\text{F}]\text{MISO}$ and PET $[^{18}\text{F}]\text{FDG}$ in two separate scanners**
 - demonstrated feasibility of multiparametric $[^{18}\text{F}]\text{MISO}/\text{FDG}$ PET-MRI
 - assessed for tumor volume, enhancement kinetics, diffusivity, and $[^{18}\text{F}]\text{FDG}/[^{18}\text{F}]\text{MISO}$ -avidity
 - descriptive statistics and voxel-by-voxel analysis of MRI and PET parameters were performed



Pinker K. et al 2016

Early CCa treatment response with mpMRI and $[^{18}\text{F}]\text{MISO}$

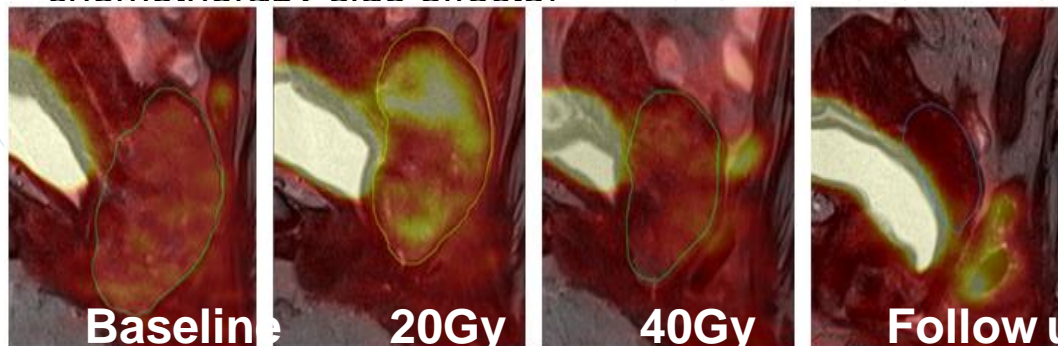
- Dataset: 6 CCa patients undergoing chemoradiotherapy, scanned with mpMRI and $[^{18}\text{F}]\text{MISO}$ PET in two separate scanners in 4 time-points
 - demonstrated feasibility of multiparametric $[^{18}\text{F}]\text{MISO}$ PET-MRI EBRT response assessment
 - high patient drop-out rate – study moved to PET/MR
 - spatio-temporal variation of hypoxia in between scanning timepoints
 - no voxel-wise correlation between hypoxia and mpMRI parameters
 - complementarity of PET and MRI derived parameters needs



Georg P. et al – manuscript in preparation

Early CCa treatment response with hybrid PET/MR using $[^{18}\text{F}]\text{MISO}$

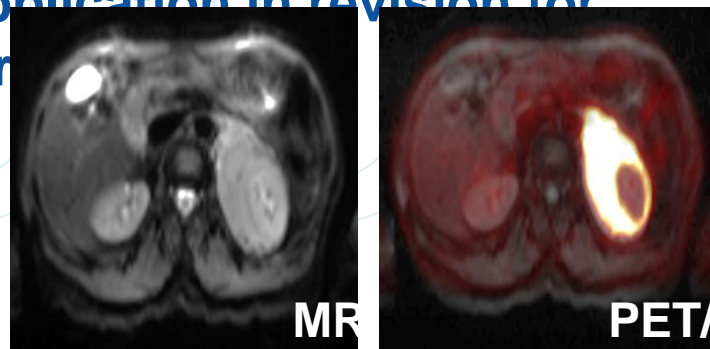
- Recruitment ongoing: 2 patients underwent full protocol, 2 patients in progress, mpMRI and $[^{18}\text{F}]\text{MISO}$ PET scans performed on a hybrid PET/MR scanner in 4 timepoints
 - ethics committee amendment approved, new SOP and imaging sequences for the PET/MR, technical developments to adjust the scanner for RO needs
 - demonstrated feasibility of multiparametric $[^{18}\text{F}]\text{MISO}$ PET/MR for EBRT response assessment (patient position as at treatment)
 - decreased patient drop-out rate (1 patient resign due to chemotherapy side effects)



Daniel M. et al - ÖGMP Jahrestagung 2016

Implementation of new PET tracers (pilot study)

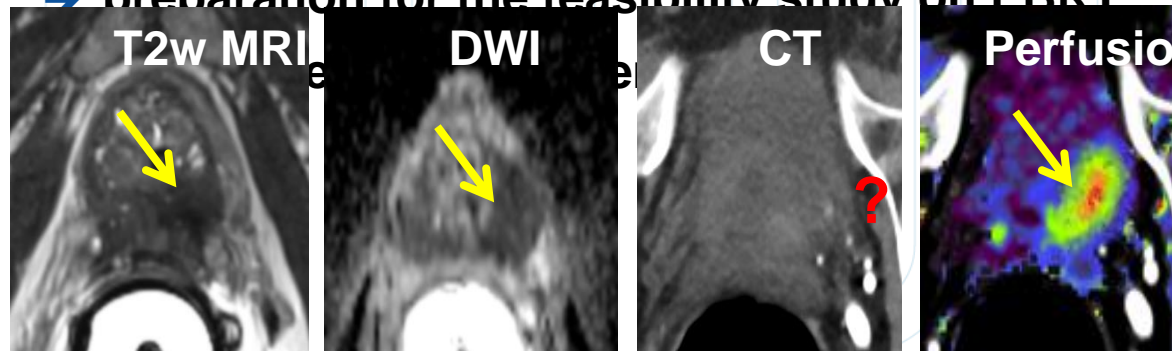
- **Early cancer treatment response using ^{68}Gd -Pentixafor (hybrid MR-PET)**
 - based on 85 studies and over 11000 patients data:
„CXCR4 over-expression is associated with poor prognosis in cancer”²⁰¹⁵
 - can be used as discriminator of necessity for more aggressive treatment
 - potential use in radiotherapy response assessment
- **Cooperation with Division of Nuclear Medicine - ethics committee application in revision for patients with cancer**
 - lung
 - pancreas
 - head and neck



Courtesy A. Haug

PCa characterization with CT perfusion (pilot study)

- **To implement CT perfusion imaging protocol in diagnosis and treatment assessment of PCa**
 - ongoing recruitment for the pre-RT baseline CT perfusion
 - optimization of the CT perfusion protocols on two diagnostic CT scanners
 - preparation for the feasibility study on EBRT

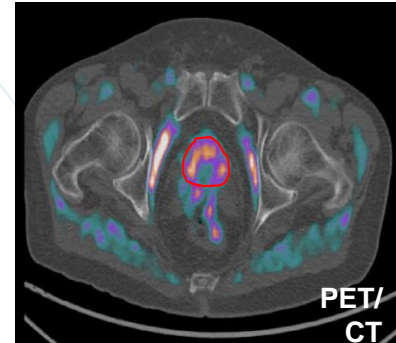


Courtesy P. Apfalter and F. Baar

Imaging data analyses

- **Based on descriptive statistics**

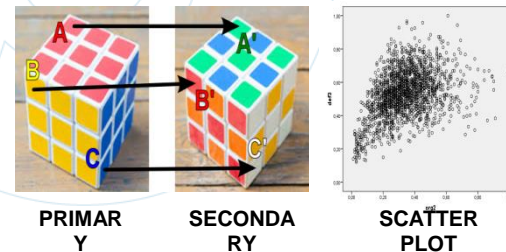
- performed in predefined regions of interest (ROIs)
- geometrical parameters (volume, distance etc.)
- grey level or biologically modeled quantitative parameters statistics (mean, max, min etc.)



Prostate NM PETresampled (PET AC)							
Role	Dataset	Method	Min	Max	Mean	Volume	Units
Planning	CTresampled (CT REFORMATTED)	HU	-948.0	3065.0	36.8	16.1cm ³	HU
**	PETresampled (PET AC)	SUV BW	1.0	3.2	1.8	16.1cm ³	g/ml

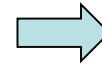
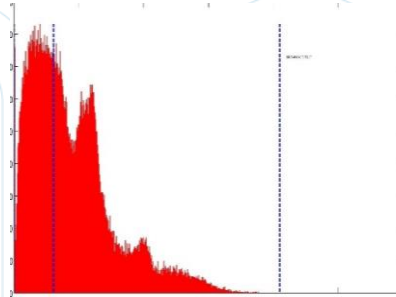
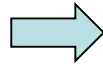
- **Based on voxel by voxel analysis**

- spatial correlation as additional degree of freedom
- requires good spatial agreement between investigated modalities (fusion or hybrid imaging)
- provides information on ROI's heterogeneity



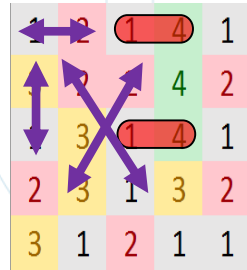
Imaging data analyses – textural features

- Based on image histogram (1st order statistics)



- Median
- Minimum
- Maximum
- Skewness
- ...

- Based on gray-level co-occurrence matrices (2nd order statistics)



	1	2	3	4
1	1	2	2	2
2	2	1	1	1
3	2	2	0	0
4	2	1	0	0



- Entropy
- Autocorrelation
- Cluster prominence
- Cluster shade
- ...

Association between pathology and texture features of mpMRI of the prostate

