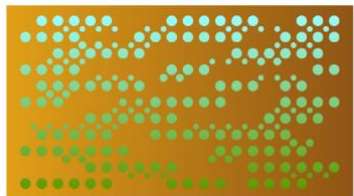


Democratizing real-time image guidance and verification: Approaches implemented on conventional linacs



DANISH CENTRE FOR PARTICLE THERAPY

Per Poulsen, Aarhus University Hospital, Denmark



Agenda

- Introduction and scope
- Real-time IGRT methods
- Real-time dose reconstruction
- Conclusion

Stereotactic Body Radiation Therapy

SBRT:

- Involves high fraction doses with steep dose gradients
- Requires high accuracy at each fraction
- Challenged by intra-fraction motion

Stereotactic Body Radiation Therapy

SBRT:

- Involves high fraction doses with steep dose gradients
- Requires high accuracy at each fraction
- Challenged by intra-fraction motion

Real-time motion adaptation:

- Could ensure high accuracy at each fraction
- Requires real-time motion monitoring

Real-time signals available on conventional linacs



Respiratory signal

kV imaging

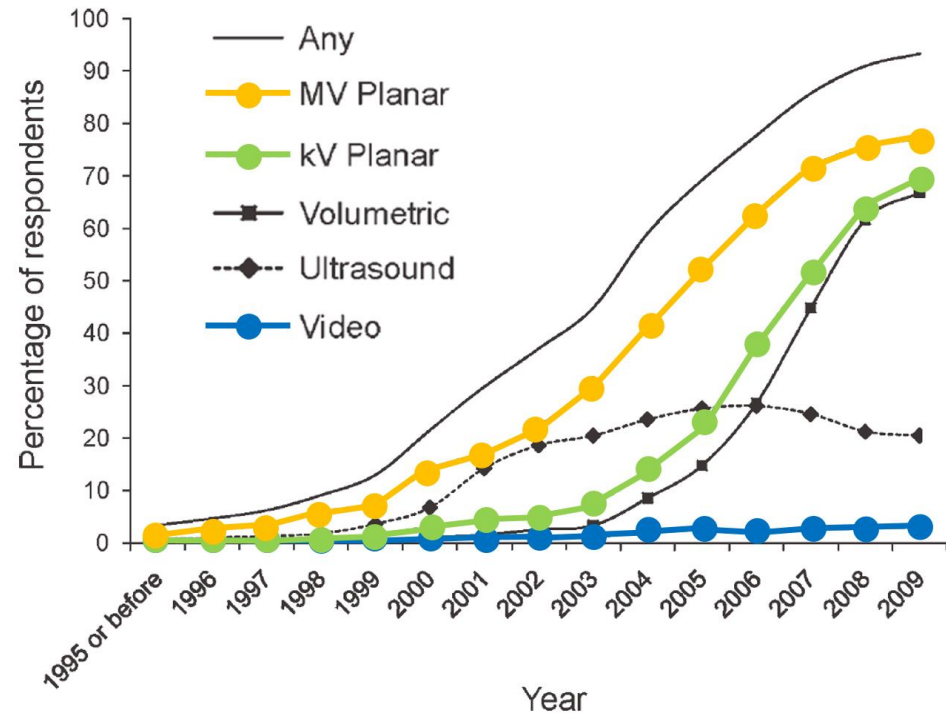
MV imaging

Adopted from Keall et al. IJROBP 2018: 102: 922-31

Surveys: Use of in-room imaging

Simpson *et al.* Cancer 2010

Survey scope: 385 MDs in US



MV imaging

kV imaging

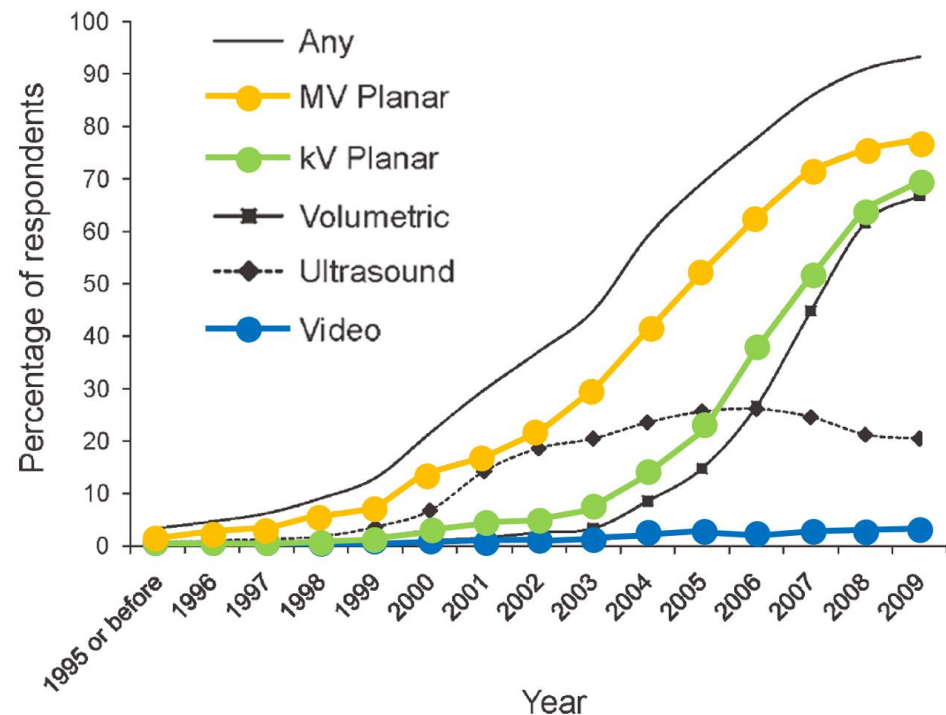
Respiratory signal



Surveys: Use of in-room imaging

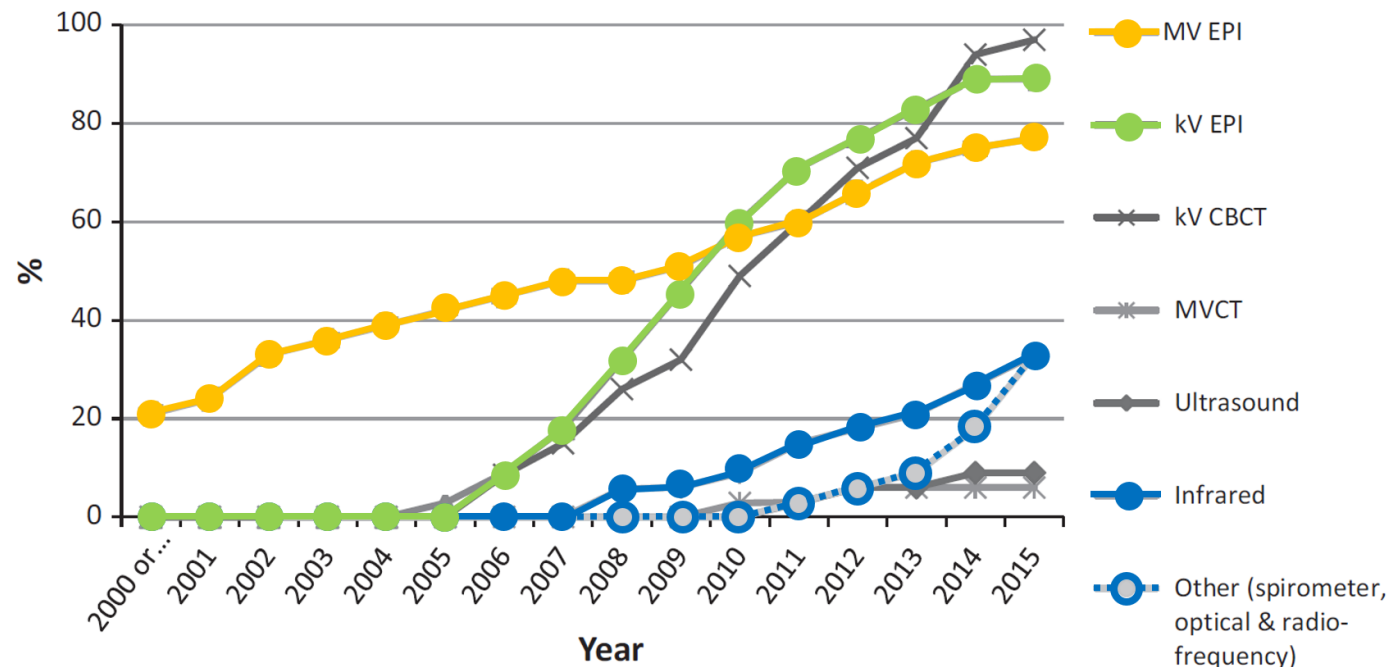
Simpson *et al.* Cancer 2010

Survey scope: 385 MDs in US



Batumalai *et al.* J Med Im Rad Onc 2017

Survey scope: 132 linacs in Australia



MV imaging

kV imaging

Respiratory signal



Real-time signals available on conventional linacs

MV imaging

- Beam's eye view
- Relatively poor contrast
- Field-of-view and time-of-view dictated by treatment plan

kV imaging

Respiratory
signal

Real-time signals available on conventional linacs

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kV imaging

- Decoupled from treatment plan
- Gives imaging dose to patient
- Perpendicular to treatment field

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- Beam's eye view
- Relatively poor contrast
- Field-of-view and time-of-view dictated by treatment plan

kV imaging

- Decoupled from treatment plan
- Gives imaging dose to patient
- Perpendicular to treatment field

Respiratory signal

- Higher frequency, lower latency
- Compatible with couch rotations
- Only external monitoring

Scope

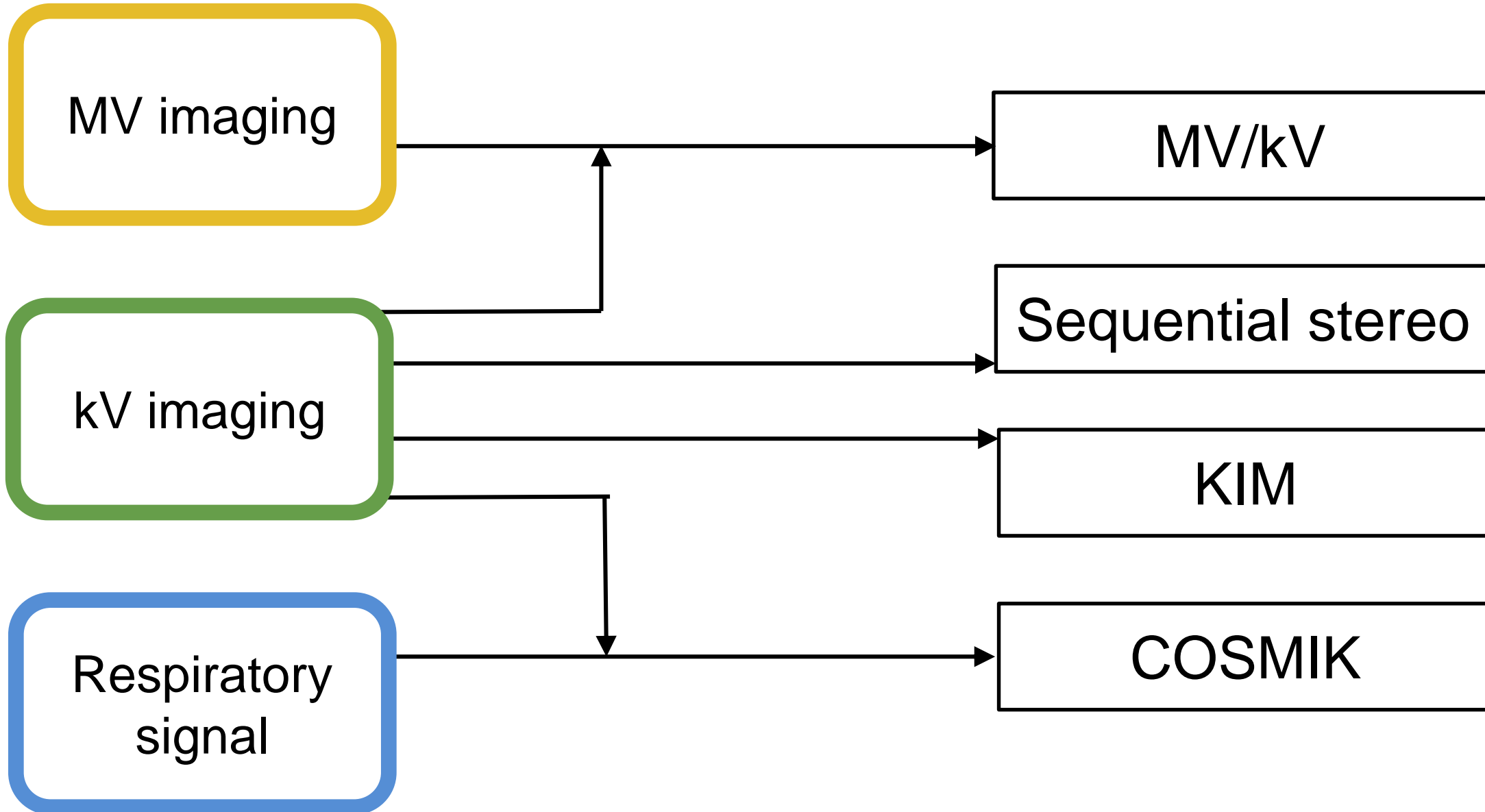
Real-time 3D image-guidance methods:

- Based on kV, MV and respiratory signals
- Used clinically during treatment delivery with conventional linacs

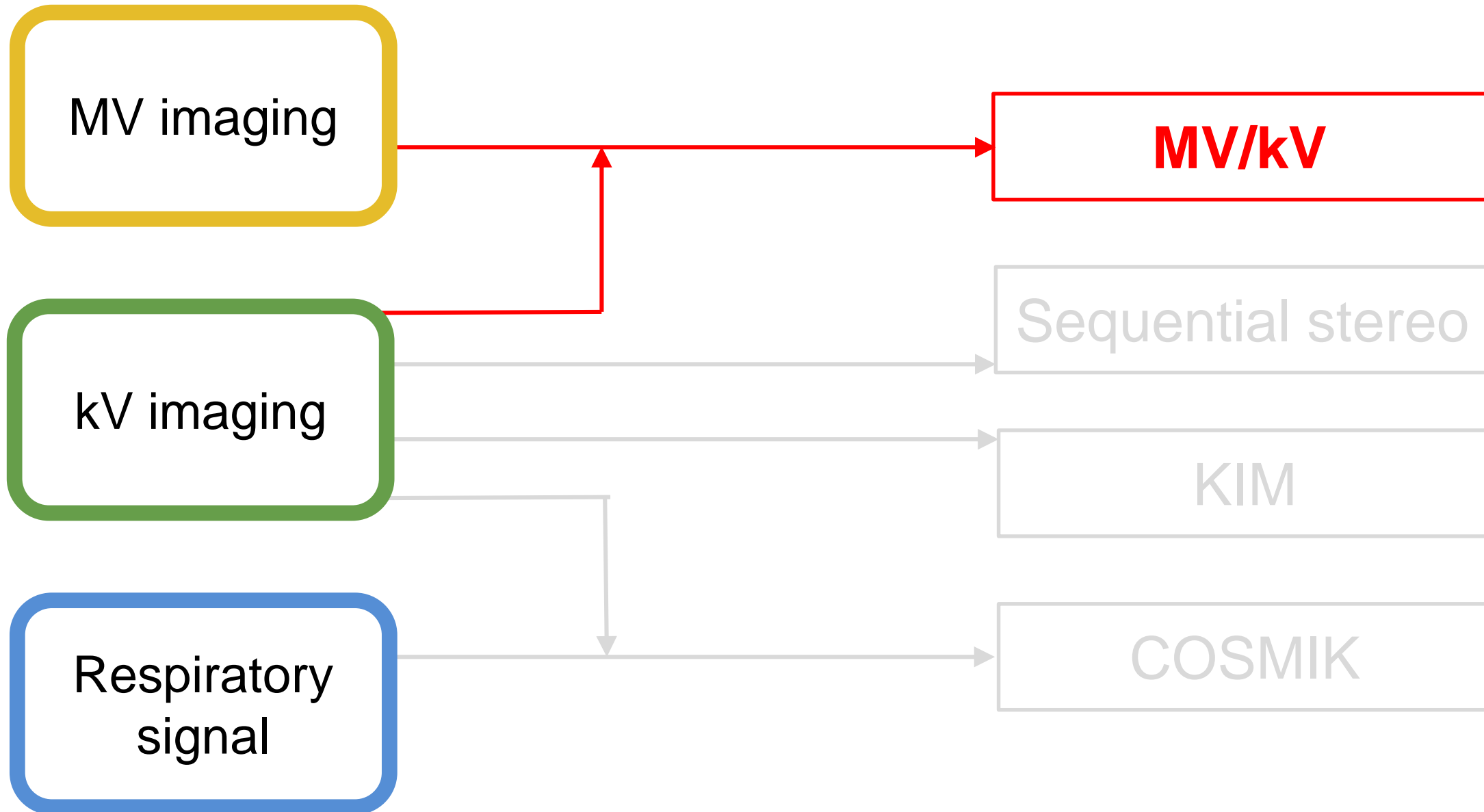
Agenda

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Four real-time IGRT methods implemented on conventional linacs



Four real-time IGRT methods implemented on conventional linacs



MV/kV imaging

VMAT prostate SBRT with implanted markers:

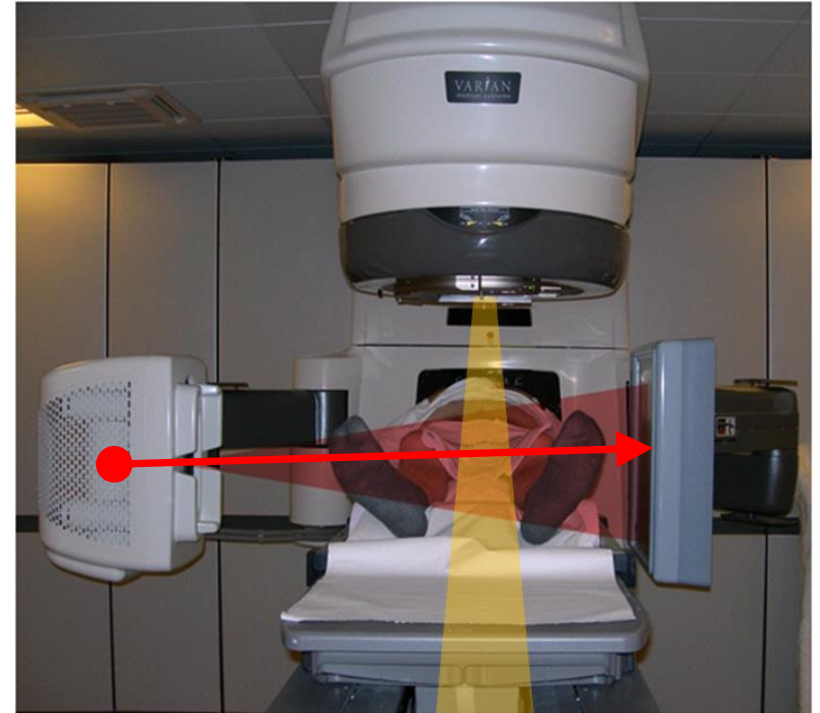
- Memorial Sloan Kettering Cancer Center

Hunt et al, J Appl Clin Med Phys 17; 473-86 (2016)

MV/kV imaging

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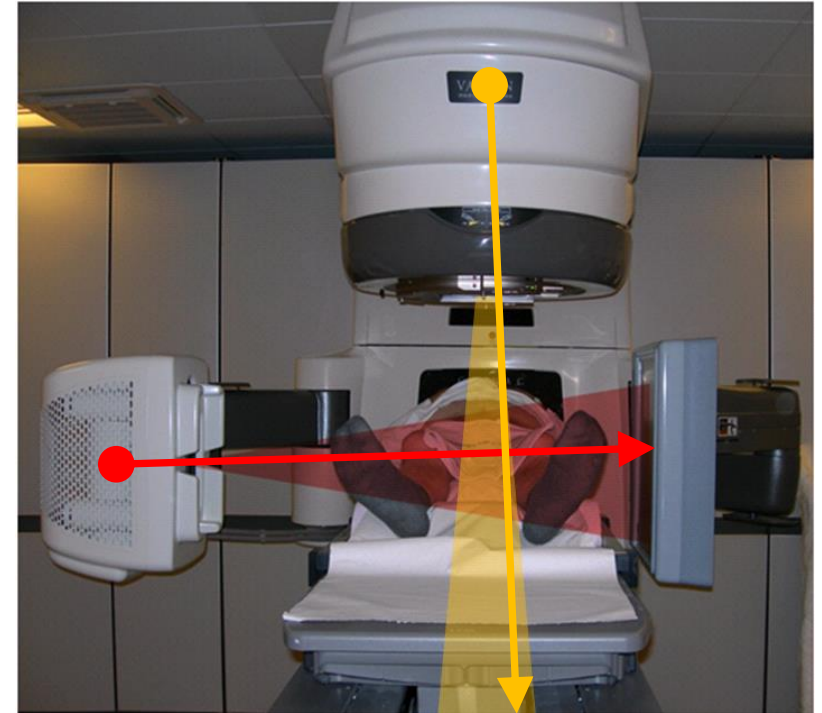


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MV/kV imaging

VMAT prostate SBRT with implanted markers:

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- Short-arc MV DTS (digital tomosynthesis, 3°) image reconstructed at kV image angles

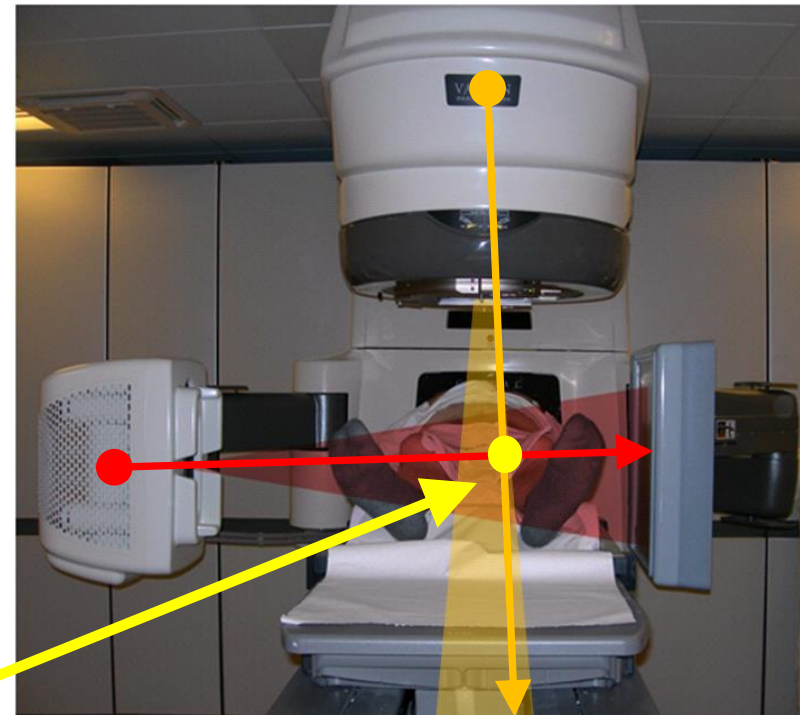


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MV/kV imaging

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- Memorial Sloan Kettering Cancer Center
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- Cine MV acquired continuously (9.5 Hz)
- Short-arc MV DTS (digital tomosynthesis, 3°) image reconstructed at kV image angles
- 3D localization by MV-kV triangulation



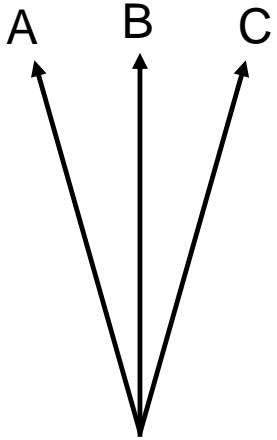
Hunt et al, J Appl Clin Med Phys 17; 473-86 (2016)

Marker visibility in MV DTS

- The markers are blocked by the MLC 60-80% of the time^(*)
- TPS script used for automatic VMAT plan manipulation
- Ensures visibility of at least one marker during the MV DTS
- Minimal degradation of plan quality

^(*) Happersett et al, J Appl Clin Med Phys 20; 120-24 (2019)

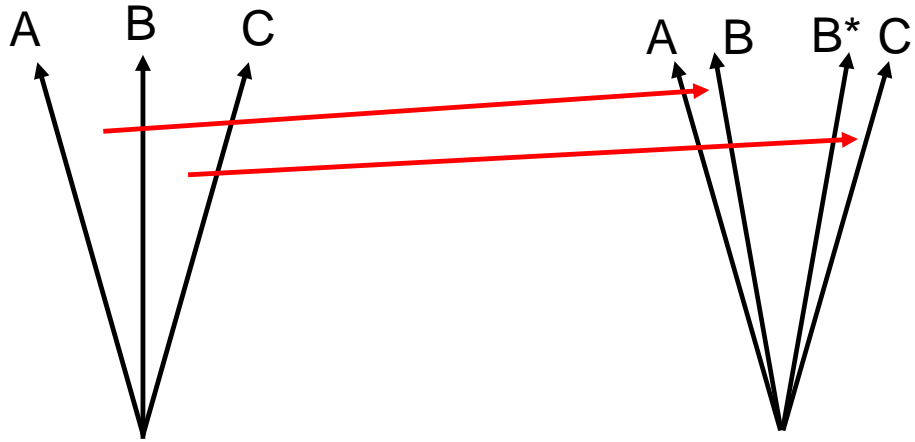
Ensuring marker visibility during MV short-arc DTS



Original control
points around
DTS position

Happersett et al, J Appl Clin Med Phys 20; 120-24 (2019)

Ensuring marker visibility during MV short-arc DTS

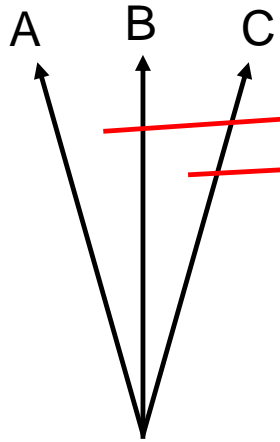


Original control
points around
DTS position

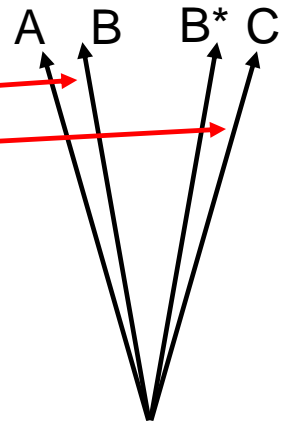
Squeeze MU
delivery to smaller
angular span

Happersett et al, J Appl Clin Med Phys 20; 120-24 (2019)

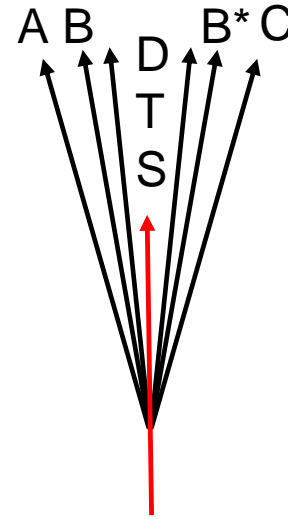
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Squeeze MU delivery to smaller angular span

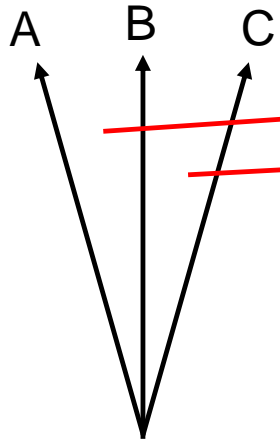


Insert DTS arc with MLC opened around at least one marker.

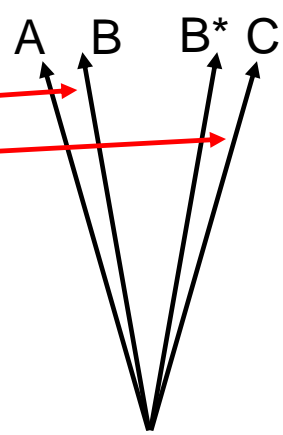
$$\text{MU}_{\text{DTS}} = 1.5\text{--}2 \text{ MU}$$

Happersett et al, J Appl Clin Med Phys 20; 120-24 (2019)

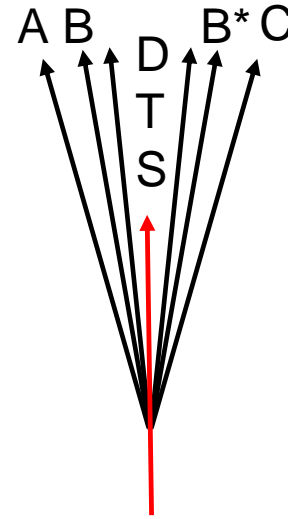
Ensuring marker visibility during MV short-arc DTS



Original control points around DTS position

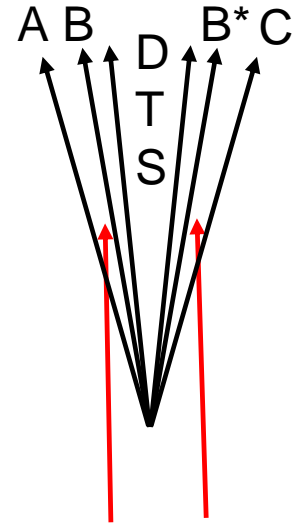


Squeeze MU delivery to smaller angular span



Insert DTS arc with MLC opened around at least one marker.

$MU_{DTS} = 1.5-2 MU$



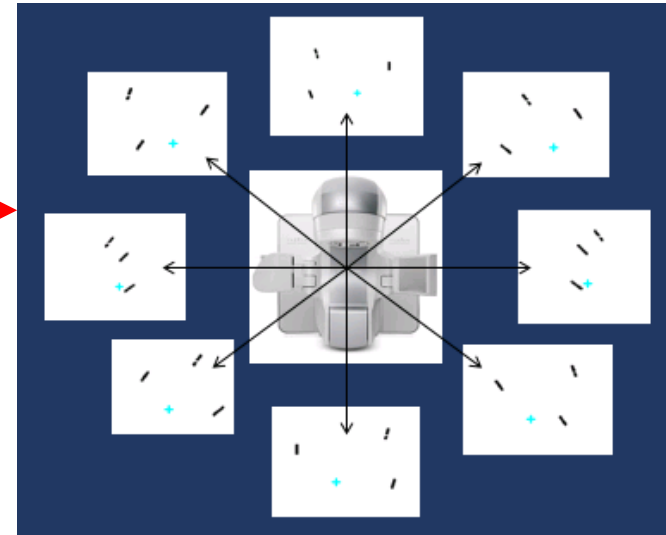
Adjust MU for MU_{DTS}

Happersett et al, J Appl Clin Med Phys 20; 120-24 (2019)

MV/kV imaging at Memorial Sloan Kettering Cancer Center

VMAT prostate SBRT implementation:

- Tracking templates created at 1° intervals 

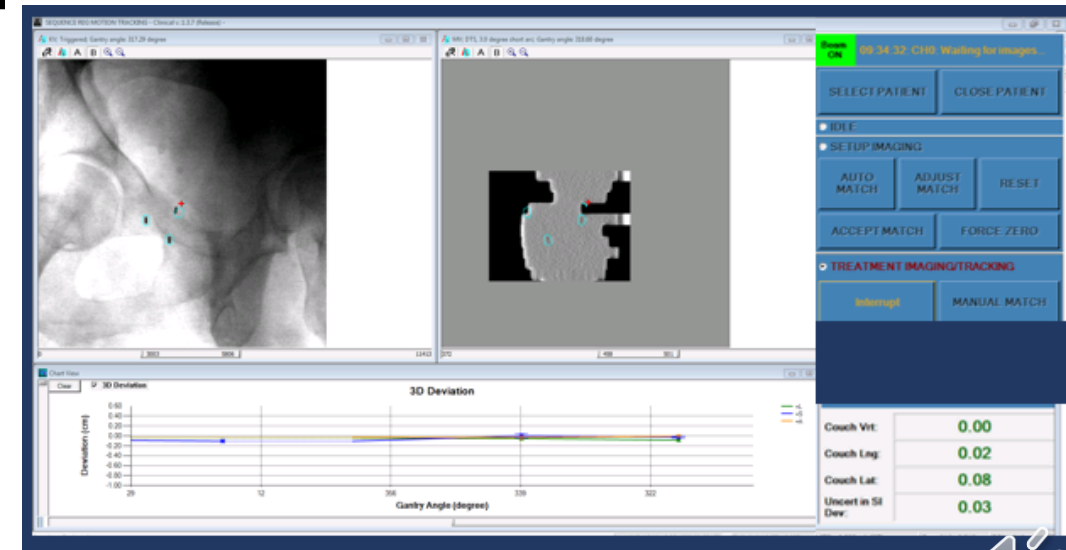
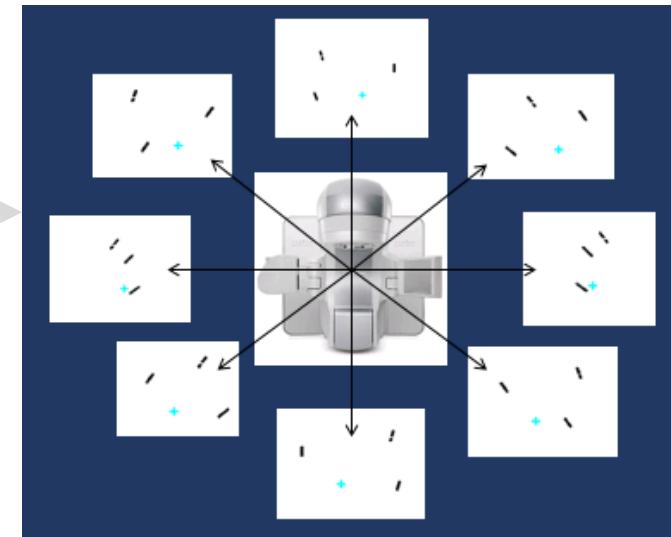


Thank you: Laura Happerset, Pengpeng Zhang, Margie Hunt, Ping Wang, Hai Pham 

MV/kV imaging at Memorial Sloan Kettering Cancer Center

VMAT prostate SBRT implementation:

- Tracking templates created at 1° intervals
- Registered with MV DTS and kV images by Sequence Reg software during treatment delivery
- Gate-off manually and correct if $>1.5\text{mm}$ error in two consecutive images



Thank you: Laura Happerset, Pengpeng Zhang, Margie Hunt, Ping Wang, Hai Pham

MV/kV imaging at Memorial Sloan Kettering Cancer Center

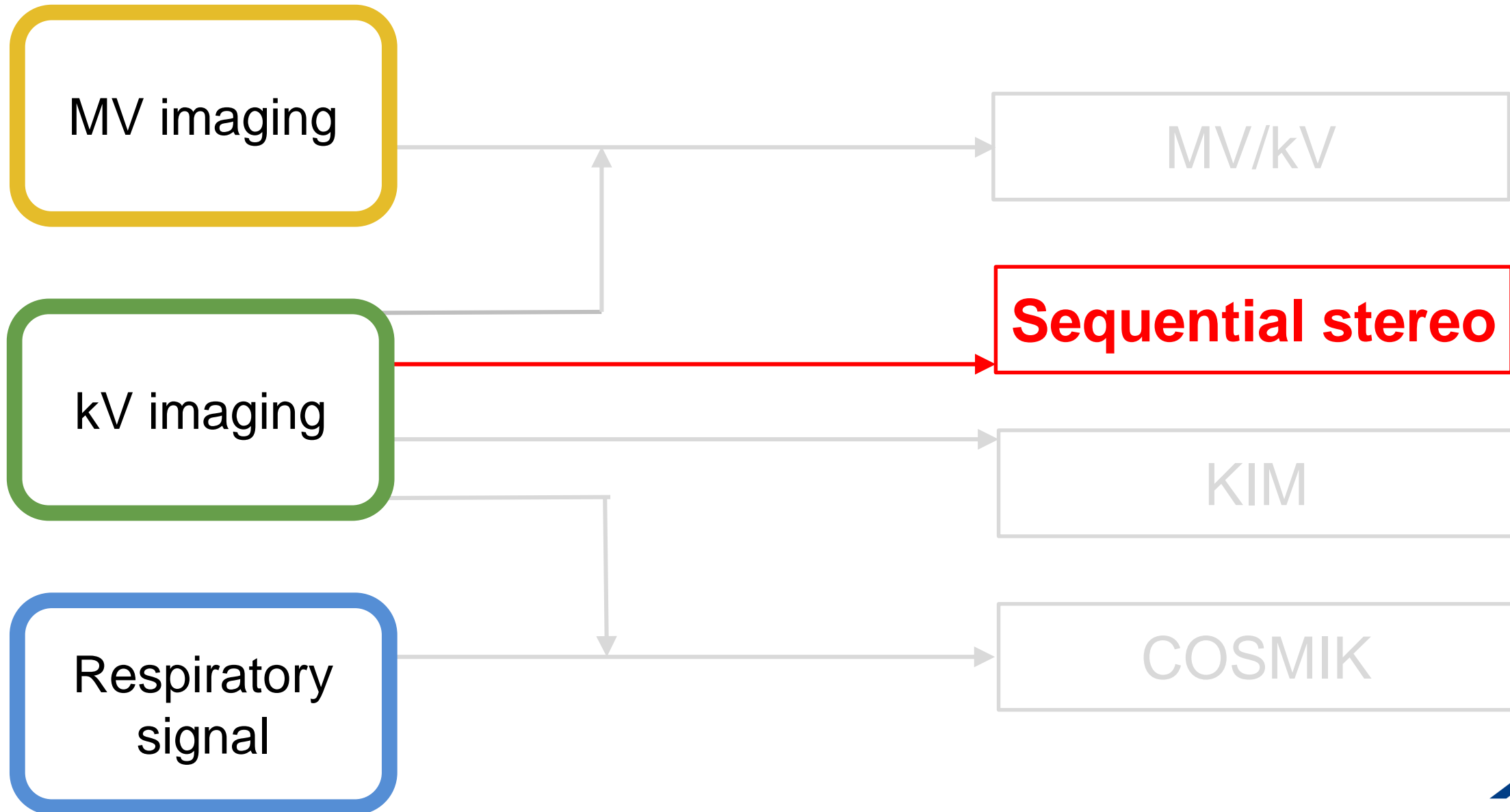
VMAT prostate SBRT application:

- 594 5-fraction prostate cancer patients treated 2016-2020
- On average 1.2 interruptions per fraction
- Prostate motion >5 mm for 10% of patients
- Median treatment time 9 minutes (measured for subset of patients)

Thank you: Laura Happerset, Pengpeng Zhang, Margie Hunt, Ping Wang, Hai Pham



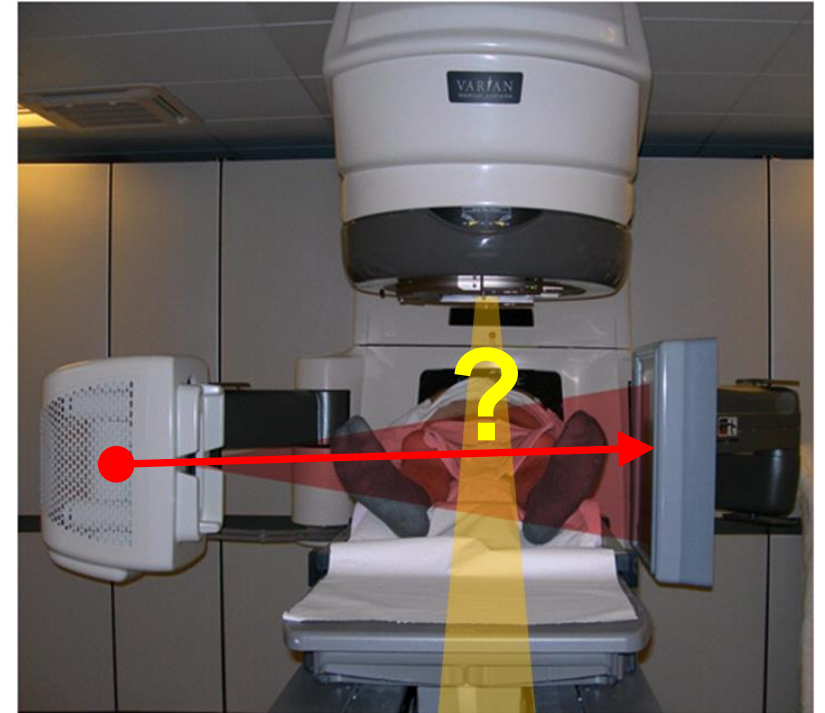
Four real-time IGRT methods implemented on conventional linacs



Sequential stereo

VMAT spine SBRT:

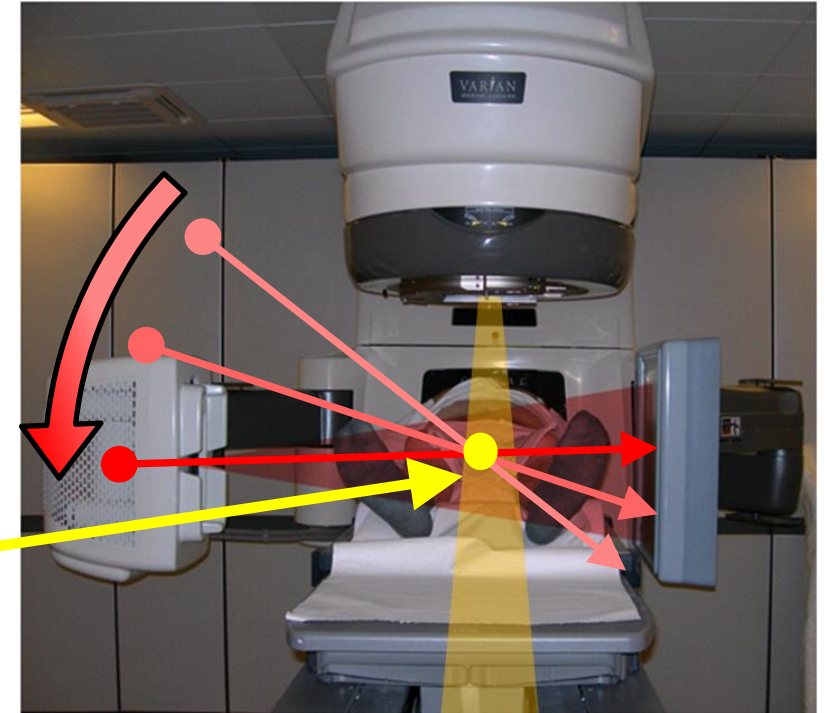
- VU University Medical Center, Amsterdam
- Continuous kV imaging (7 Hz)
- Match on spine with DRRs from planning CT



Sequential stereo

VMAT spine SBRT:

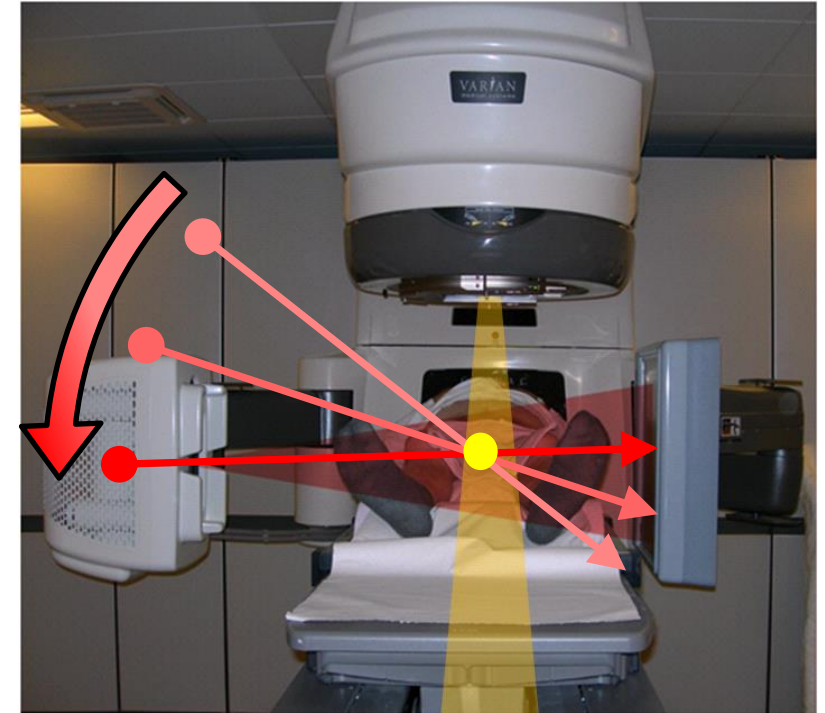
- VU University Medical Center, Amsterdam
- Continuous kV imaging (7 Hz)
- Match on spine with DRRs from planning CT
- **Triangulation with 2-8 previous kV images**
- **Assume no motion along current ray line since the previous images**



Sequential stereo

Selection of previous kV images for triangulation:

- acquired $14-72^\circ$ prior to current image
- ray line $<1\text{mm}$ from current ray line
- all 2-8 selected ray lines intersect in a small volume close to current ray line



Sequential stereo at VU University Medical Center, Amsterdam

VMAT spine SBRT implementation:

- Tracking templates (DRRs) created at 1° intervals
- Registered with streamed kV images during treatment delivery
- Gate-off manually and correct (by CBCT) if >1 mm error in any direction



Sequential stereo at VU University Medical Center, Amsterdam

First clinical online real-time experiences^(*):

- 10 spine SBRT fractions of 3 patients:
- Images analyzed at ~1Hz (limited by computing speed)
- 2 beam interruptions in total due to >1mm errors

^(*) Hazelaar et al, IJROBP 2018



Sequential stereo at VU University Medical Center, Amsterdam

First clinical online real-time experiences^(*):

- 10 spine SBRT fractions of 3 patients:
- Images analyzed at ~1Hz (limited by computing speed)
- 2 beam interruptions in total due to >1mm errors

Further clinical experiences with real-time sequential stereo^(**):

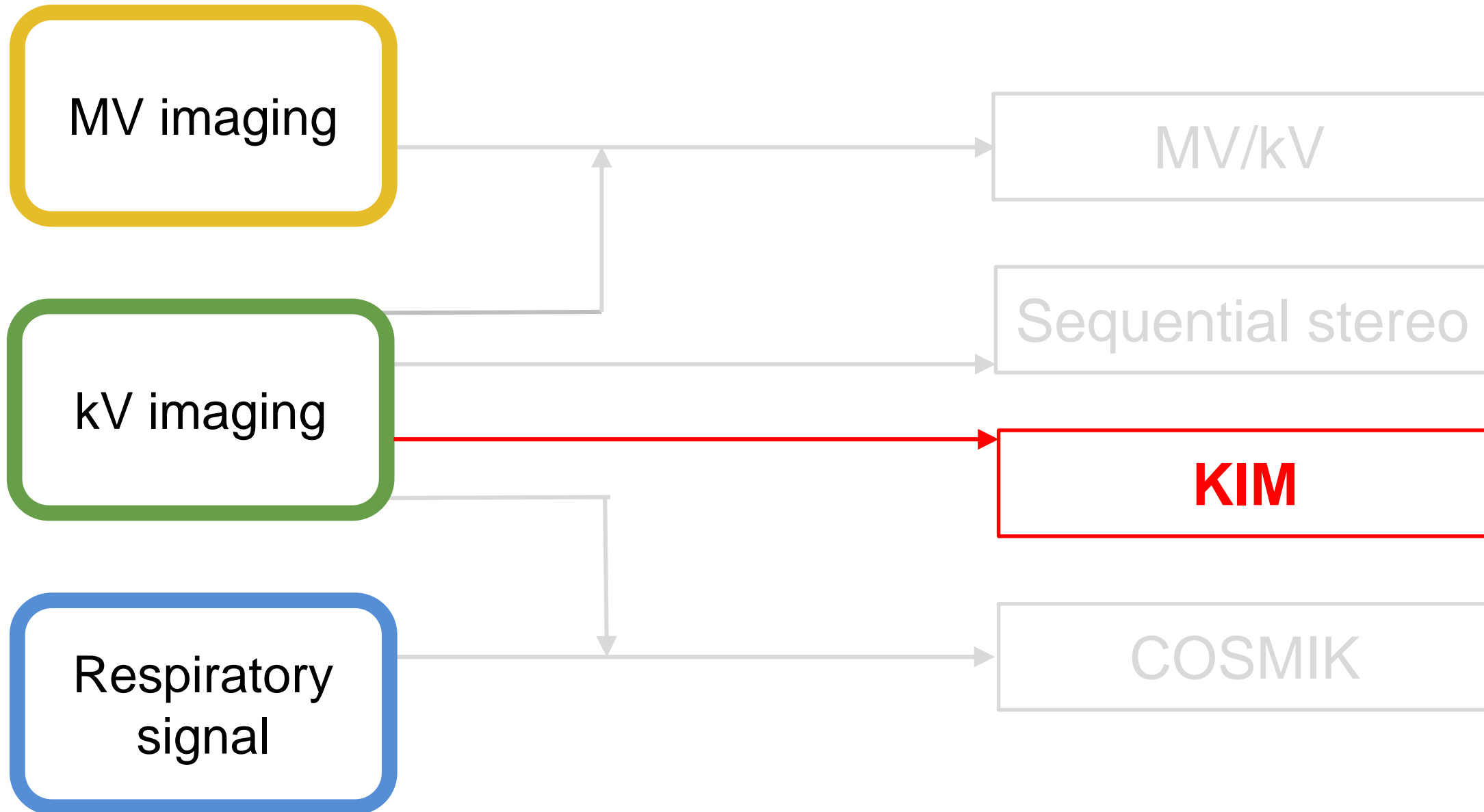
- ~40 spine SBRT
- DIBH lung SBRT
- Airway tracking (proximal bronchial tree) for central lung SBRT

^(*) Hazelaar et al. IJROBP 2018

^(**) W Verbakel, private communication



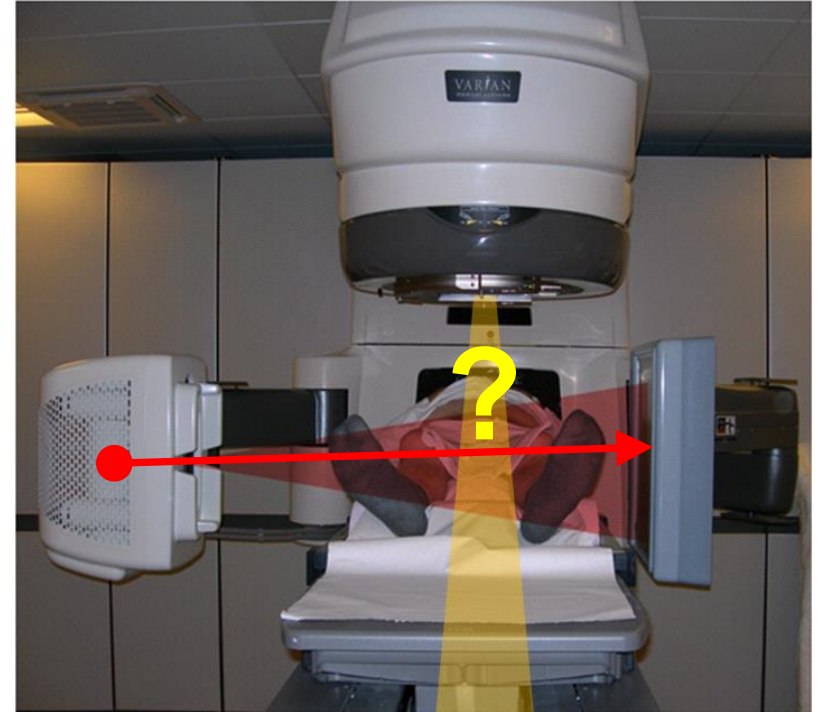
Four real-time IGRT methods implemented on conventional linacs



KIM (Kilovoltage Intrafraction Monitoring)

KIM:

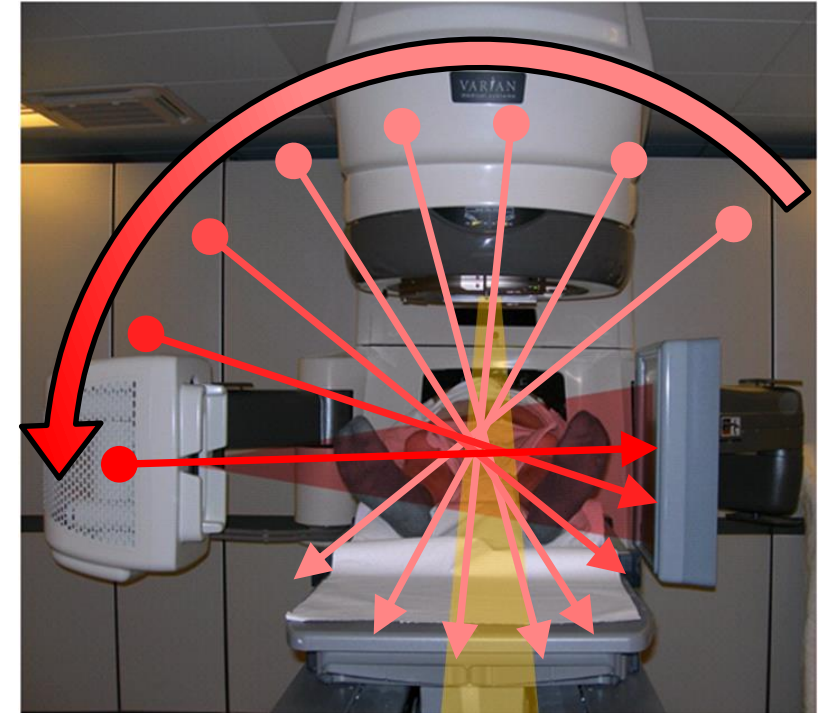
- Continuous kV imaging (5-11 Hz)



KIM (Kilovoltage Intrafraction Monitoring)

KIM:

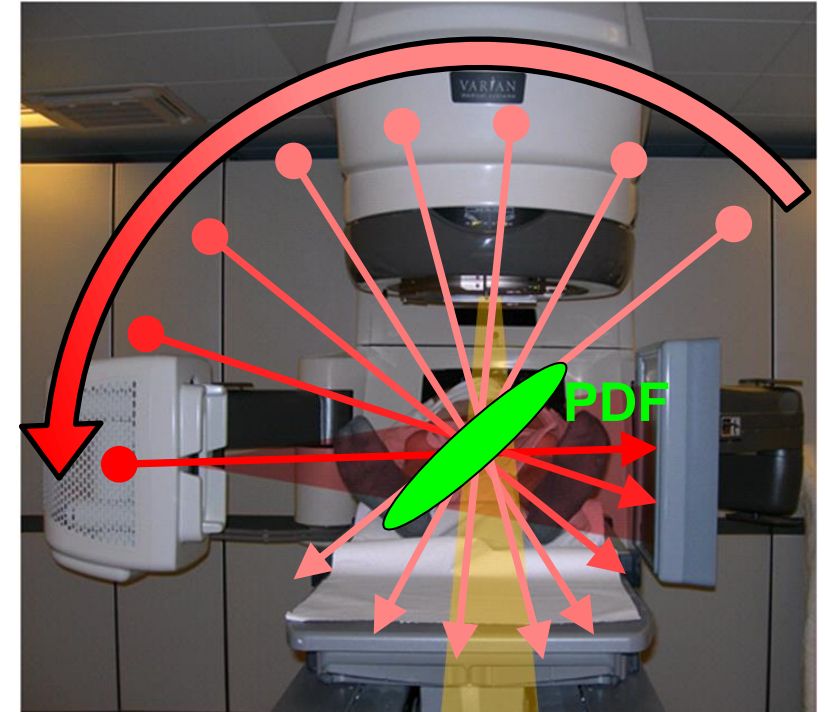
- Continuous kV imaging (5-11 Hz)
- Use all previous images in $>120^\circ$ angular span



KIM (Kilovoltage Intrafraction Monitoring)

KIM:

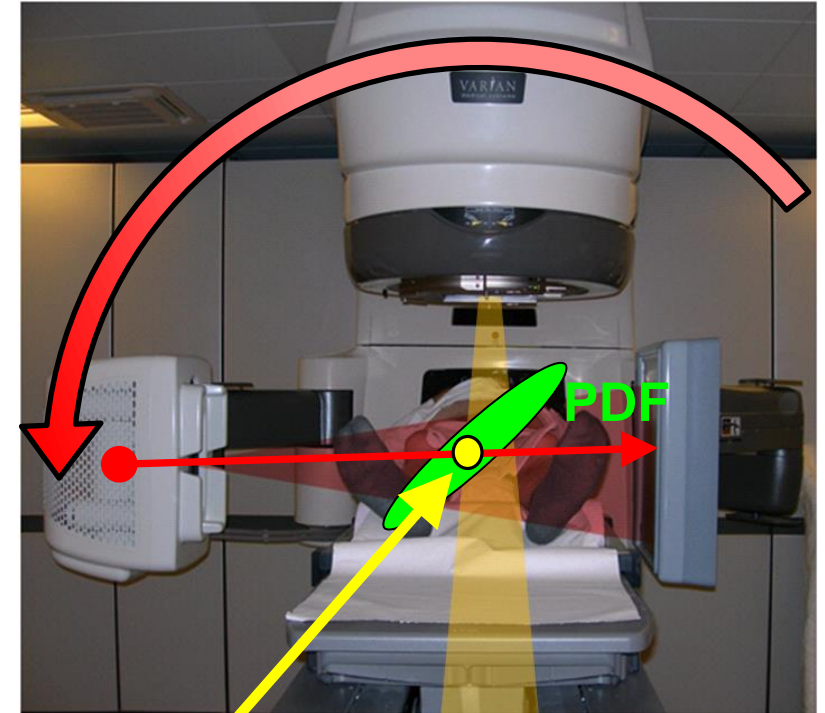
- Continuous kV imaging (5-11 Hz)
- Use all previous images in $>120^\circ$ angular span
- Find the 3D Gaussian Probability Density Function (PDF) that best describes the target motion (by maximum likelihood estimation)



KIM (Kilovoltage Intrafraction Monitoring)

KIM:

- Continuous kV imaging (5-11 Hz)
- Use all previous images in $>120^\circ$ angular span
- Find the 3D Gaussian Probability Density Function (PDF) that best describes the target motion (by maximum likelihood estimation)
- Find the most likely target position along the ray line

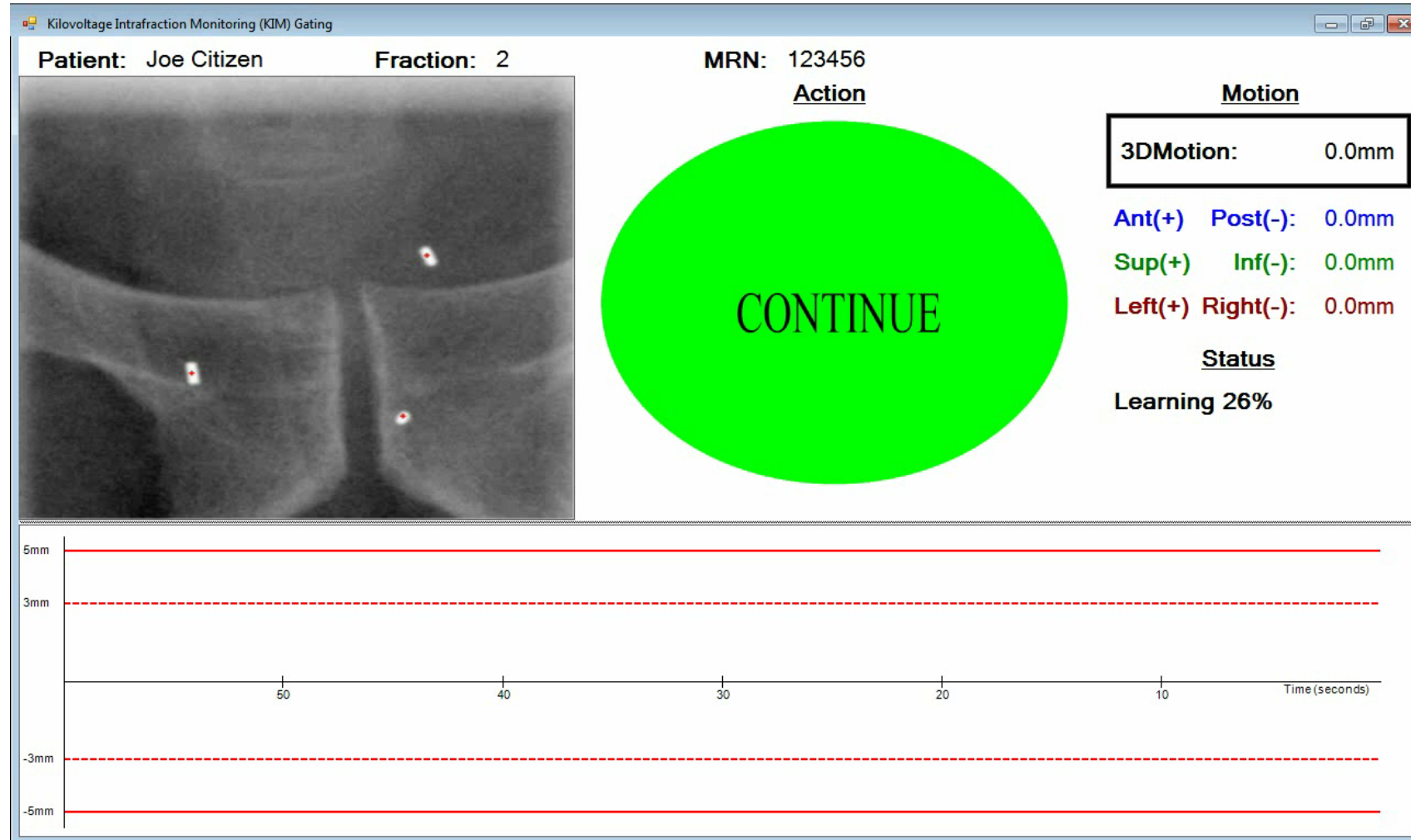


Online real-time KIM

Online real-time application:

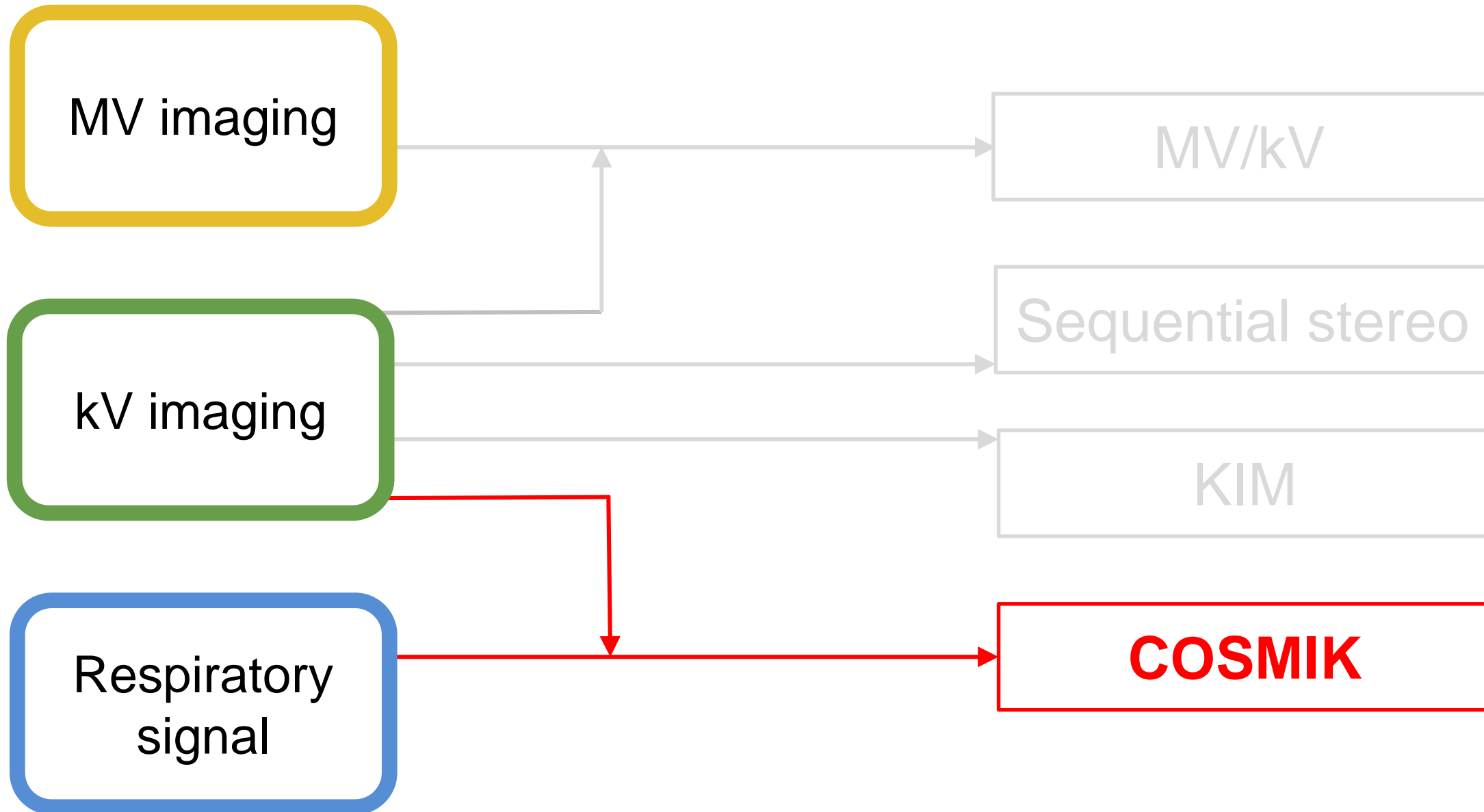
- Northern Sydney Cancer Center and four other Australian clinics
- Implemented on Varian Clinac, Varian TrueBeam, Elekta Synergy linacs
- 120 prostate cancer patients:
 - Gate-off and correct couch if $>3\text{mm}$ error in $>5\text{s}$ (~2000 fractions)
 - MLC tracking (49 fractions)
- 2 liver SBRT patients treated in breath-hold (7 fractions)

Prostate KIM at Northern Sydney Cancer Center



Video from Paul Keall, Doan Trang Nguyen, Jeremy Booth

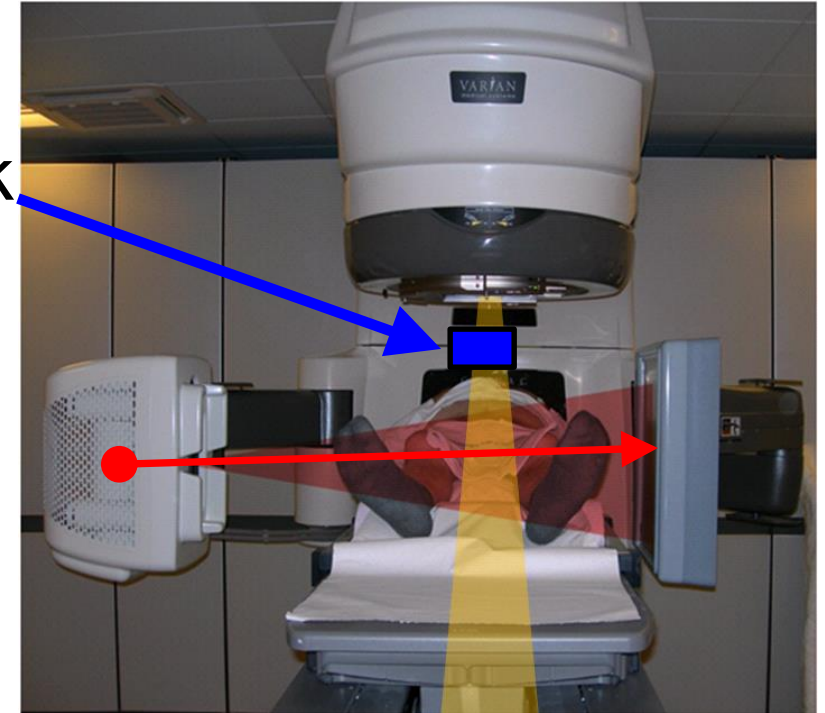
Four real-time IGRT methods implemented on conventional linacs



COSMIK (Combined Optical and Sparse Monoscopic Imaging with KV x-rays)

COSMIK:

- Continuous monitoring of external marker block
- Sparse kV imaging of implanted markers (every 3 sec)
- Only for respiratory motion



COSMIK workflow (liver SBRT)

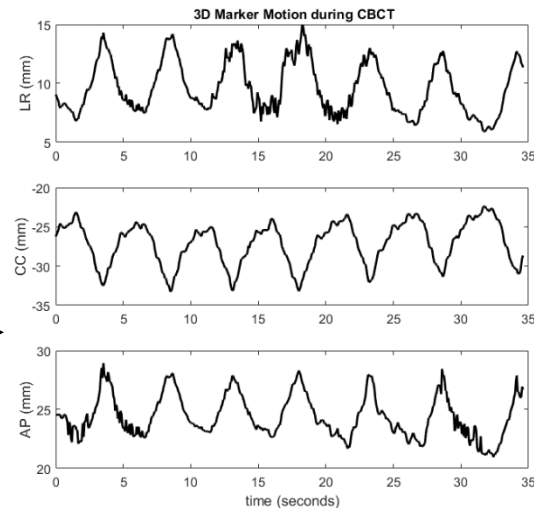
Pre-treatment setup CBCT:

1. Extract internal 3D marker trajectory using KIM
2. Establish correlation model between external and internal motion



CBCT projections (liver SBRT)

KIM



3D marker motion

External motion

External-internal
motion correlation
model

COSMIK workflow (liver SBRT)

Pre-treatment setup CBCT:

1. Extract internal 3D marker trajectory using KIM
2. Establish correlation model between external and internal motion

During treatment delivery:

- Continuous external optical monitoring (20 Hz):
 - 3D internal marker motion estimated from correlation model
- Sparse kV imaging (0.33 Hz):
 - 3D internal marker position estimated with KIM
 - Update correlation model to account for tumor drift

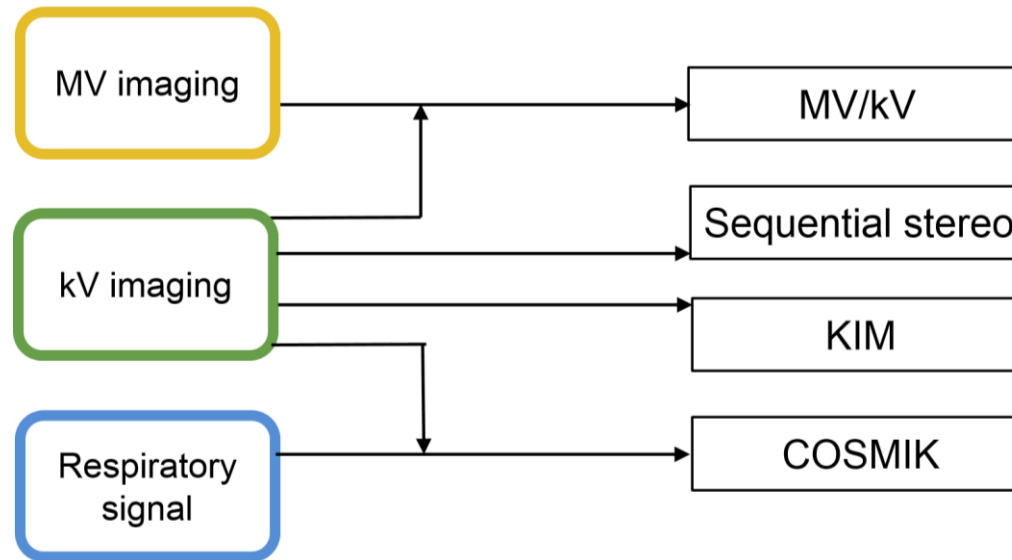
COSMIK at Aarhus University Hospital

Online real-time application:

- ~20 liver SBRT patients
- 1 lung cancer patient (markers in mediastinal lymph nodes)


- Real-time motion-including tumor dose reconstruction

Common characteristics of all four real-time IGRT methods



- Implemented clinically on conventional linacs
- Research software, not commercially available
- (Sub)-millimeter accuracy
- 3 of 4 methods currently rely on implanted markers

Summary of clinically implemented 3D real-time IGRT methods

	MV/kV	Sequential stereo	KIM	COSMIK
Principle	MV short-arc DTS Triangulation	Triangulation	Monoscopic imaging	Monoscopic imaging Breathing correlation
Tumor sites	Prostate	Spine	Prostate, liver	Liver, mediastinal LN
Treatment adaptation	Gate-off and adjust couch	Gate-off and adjust couch	Gate-off and adjust couch. MLC tracking	None (only real-time dose reconstruction)
Markers	Yes	No	Yes	Yes
Non-coplanar	No	No	No	Yes
Frequency	0.1-0.2 Hz (every 20°)	1 Hz (7Hz imaging)	5-11 Hz	20 Hz
Motion type	Small	Small (or periodic)	Any	Respiratory motion
Comments	<ul style="list-style-type: none"> • Low kV dose • Prior imaging not needed • Requires marker in MLC aperture 	<ul style="list-style-type: none"> • Requires prior images with same target position 	<ul style="list-style-type: none"> • Requires learning • Requires stable motion PDF 	<ul style="list-style-type: none"> • Requires learning • Low kV dose • Continuous monitoring at Fx • Low latency 

Agenda

- Introduction
- Real-time IGRT methods
- Real-time dose reconstruction
- Conclusion

Real-time motion-including dose reconstruction

Motivation:

- Dose errors from motion >> Dose errors from machine QA

Dose reconstruction during treatment delivery:

- Enabled by real-time IGRT
- Important tool for real-time verification

Real-time motion-including dose reconstruction

DoseTracker in-house software:

- Very simple dose calculation
 - Water density
 - Flat patient surface
 - Same phantom scatter in all depths

Real-time motion-including dose reconstruction

DoseTracker in-house software:

- Very simple dose calculation
 - Water density
 - Flat patient surface
 - Same phantom scatter in all depths
-but flexible and fast
 - Any set of calculation points
 - ~100ms for 20,000 calculation points in a tumor
 - **Includes motion**

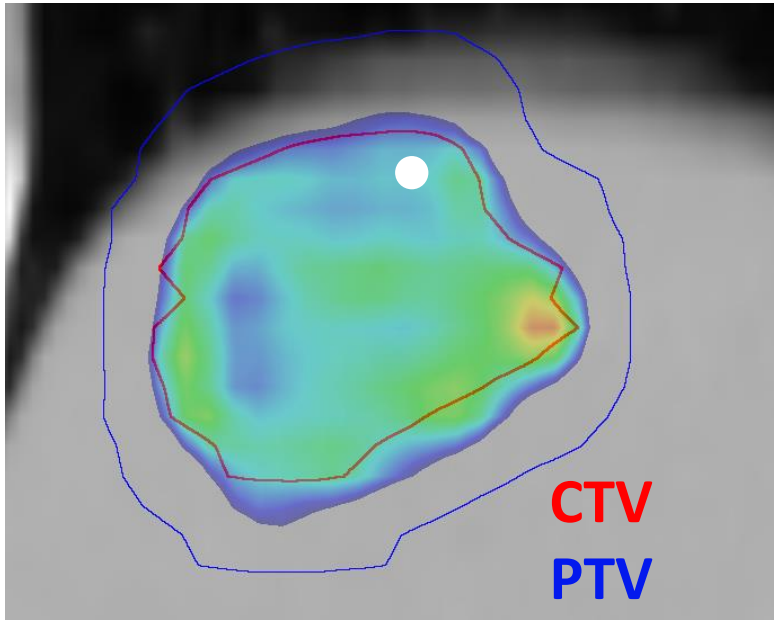


Real-time dose reconstruction

First online application:

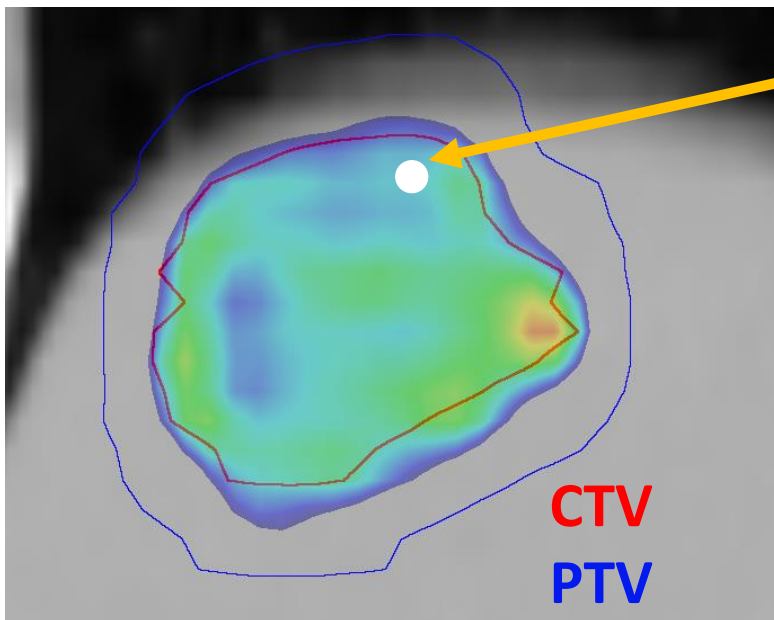
- Aarhus University Hospital
- 7 VMAT liver SBRT patients (10 fractions)
- Tumor position (COSMIK) and linac parameters streamed to DoseTracker
- Dose reconstructed in the PTV (1700-4500 calculation points, ~9 Hz)

Example: Dose reconstruction in a single point



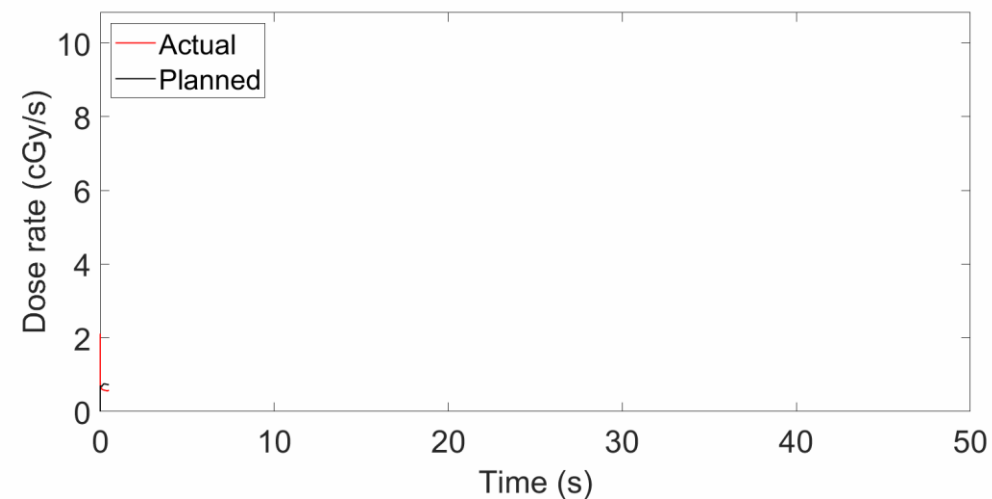
Planned dose to liver tumor
(95-107% shown)

Example: Dose reconstruction in a single point

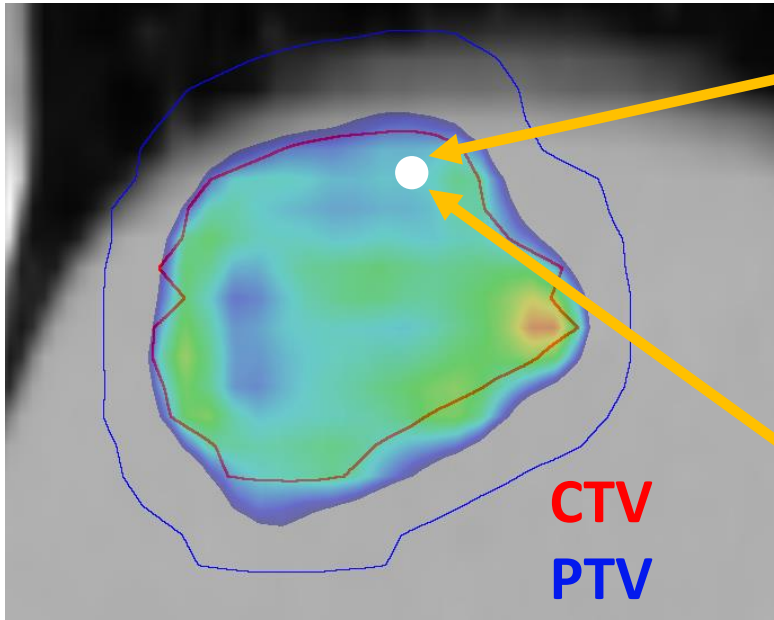


Planned dose to liver tumor
(95-107% shown)

DoseTracker (during treatment)

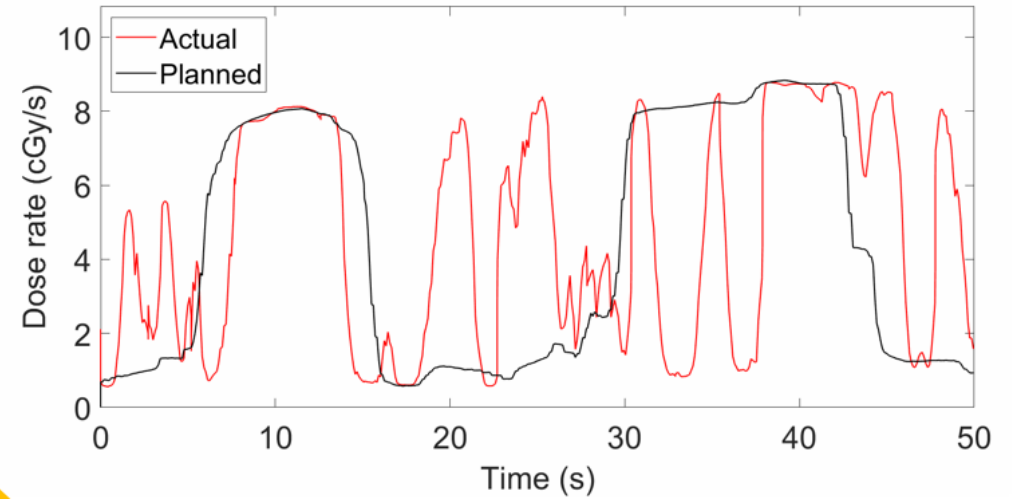


Example: Dose reconstruction in a single point

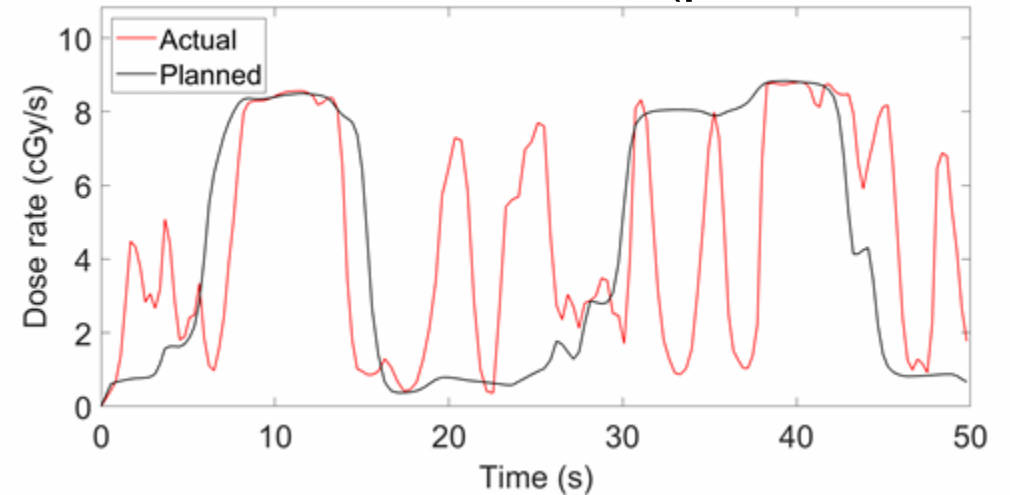


Planned dose to liver tumor
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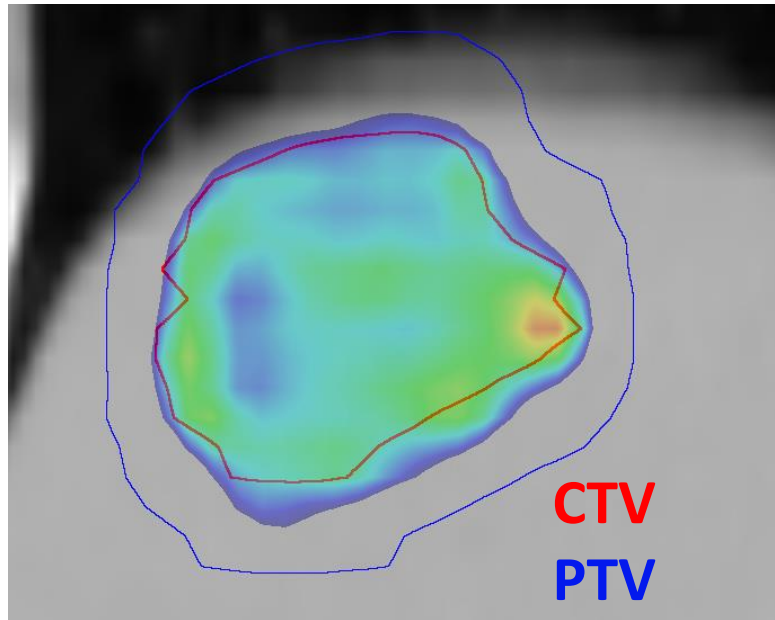
DoseTracker (during treatment)



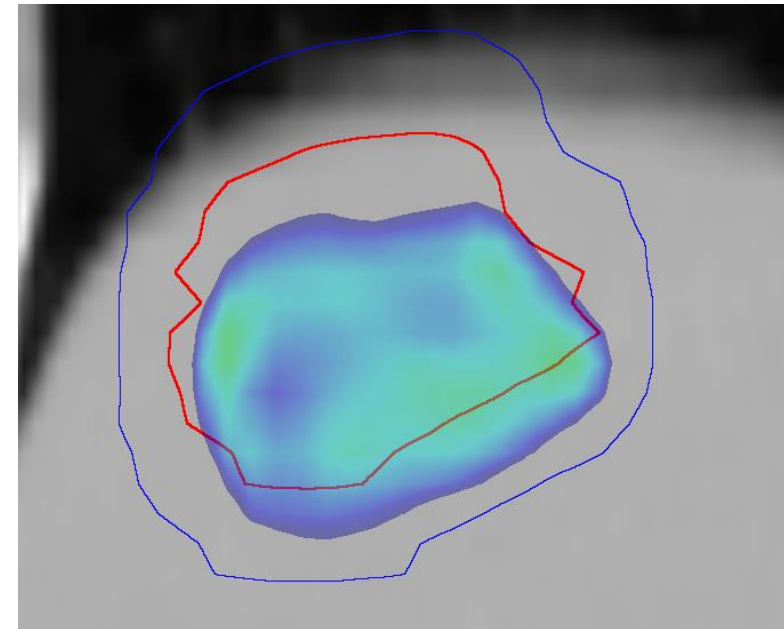
TPS, isocenter shift (post-treatment)



Planned and delivered dose distribution

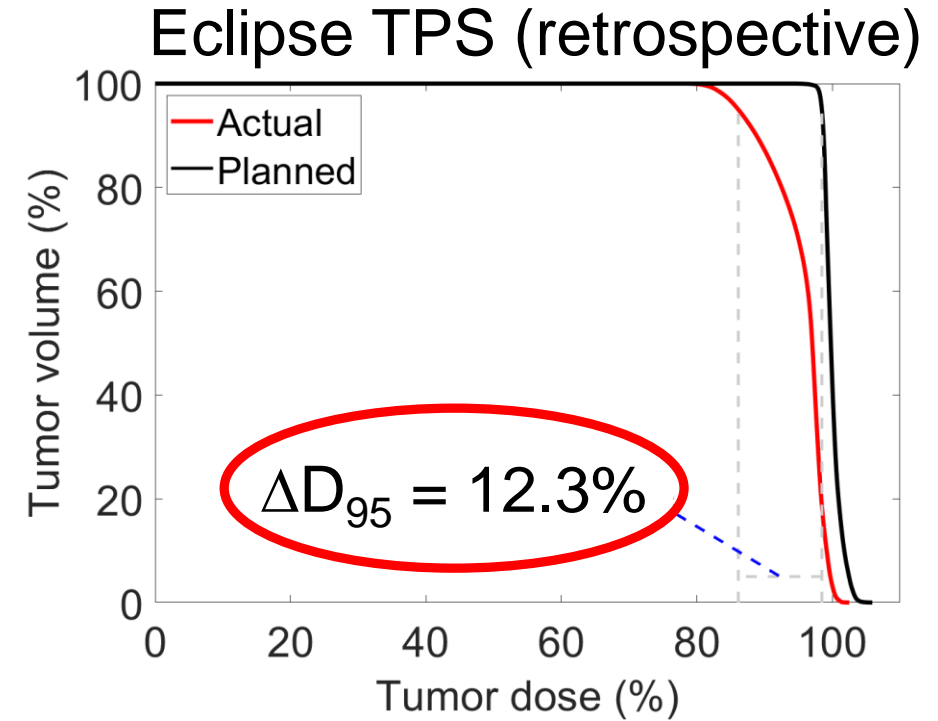
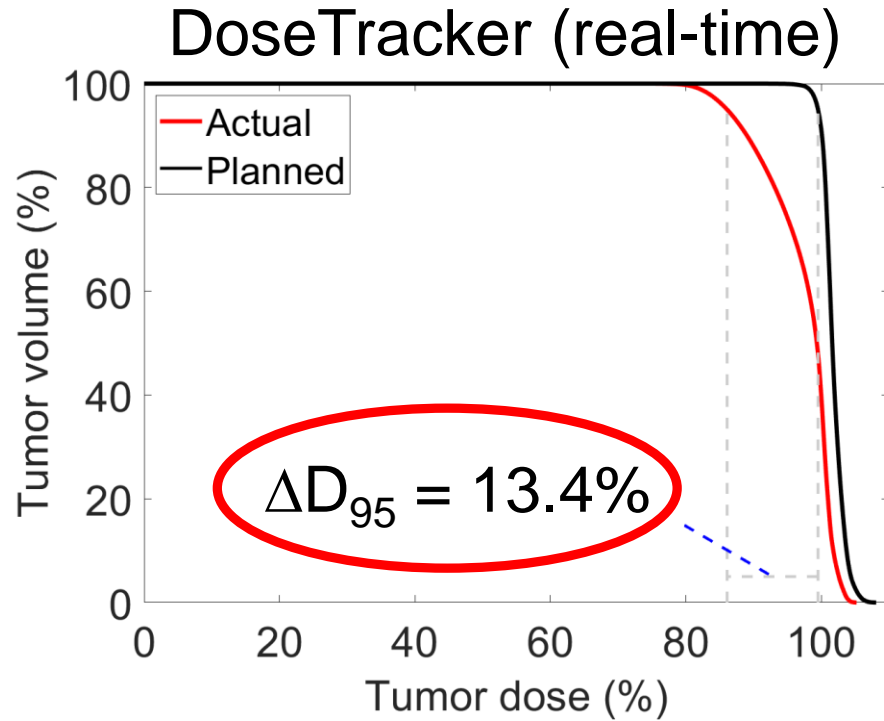


Planned dose to liver tumor
(95-107% shown)

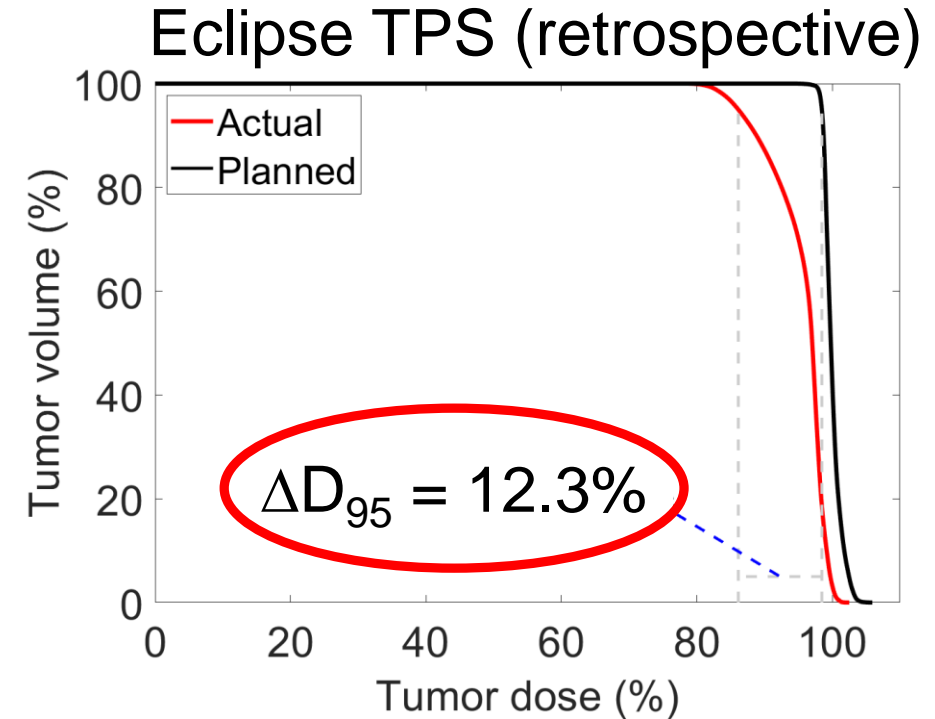
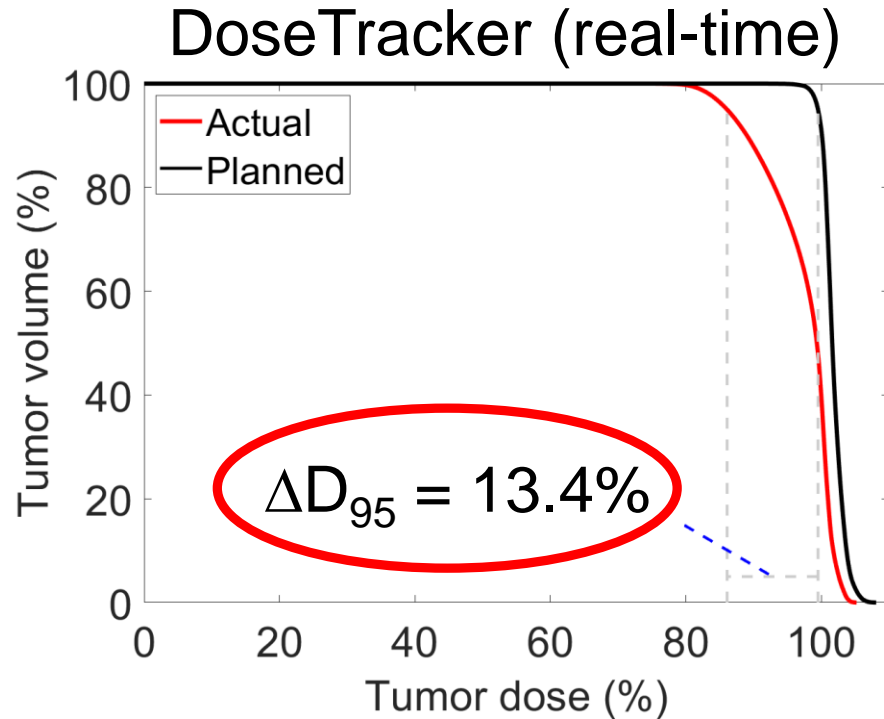


Delivered dose
(95-107% shown)

Motion-induced reduction in CTV D_{95}



Motion-induced reduction in CTV D_{95}



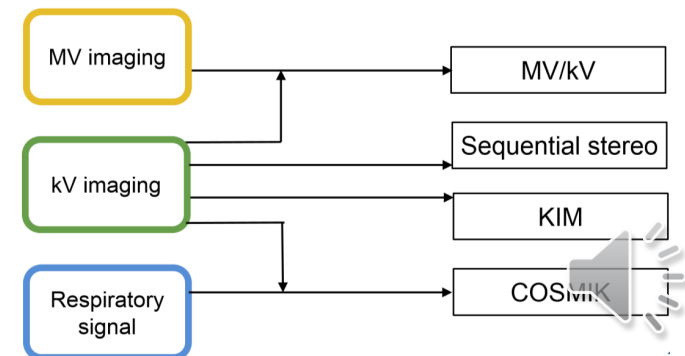
RMS error in real-time calculated ΔD_{95} for all 10 fractions: 1.3%-points

Agenda

- Introduction
- Real-time IGRT methods
- Real-time dose reconstruction
- Conclusion

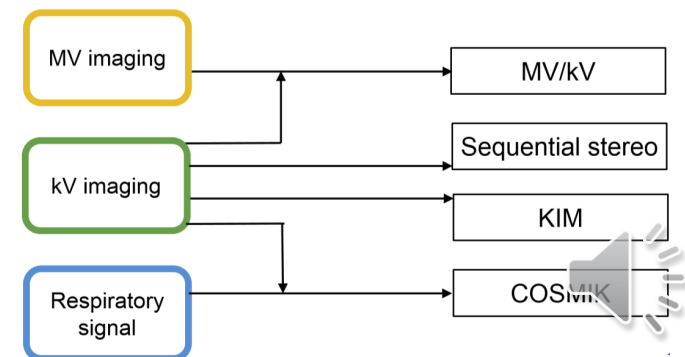
Conclusions: Real-time IGRT on conventional linacs

- Technologies are being developed by researchers
- Used for real-time treatment adaptation or tumor dose reconstruction
- Potential for widespread use



Conclusions: Real-time IGRT on conventional linacs

- Technologies are being developed by researchers
- Used for real-time treatment adaptation or tumor dose reconstruction
- Potential for widespread use
- Broad adoption requires commercial solutions closely integrated with the clinical workflow
- MR-linacs may become driving a force for broader adoption
- IMRT, VMAT, CBCT: Fast clinical implementation of new technology
- Further development of markerless localization and image dose reduction may facilitate broader adoption of real-time IGRT.



Acknowledgements

Aarhus University Hospital:

- Thomas Ravkilde
- Simon Skouboe
- Esben Worm
- Morten Høyer
- Britta Weber

VU University Medical Center:

- Wilko Verbakel

Bern University Hospital:

- Jenny Bertholet

Varian Medical Systems

University of Sydney:

- Paul Keall
- Doan Trang Nguyen
- Jeremy Booth
- Ricky O'Brien

Memorial Sloan Kettering Cancer Center:

- Pengpeng Zhang,
- Laura Happersett
- Margie Hunt
- Ping Wang
- Hai Pham