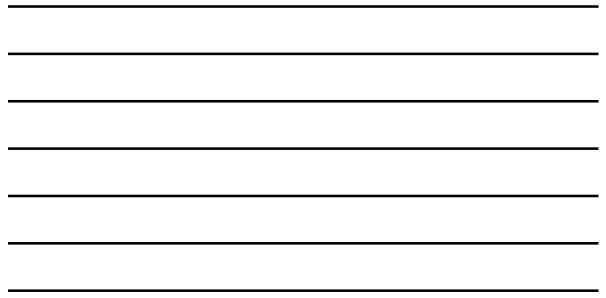


Real-time Respiratory Tumor Tracking with Implanted Markers

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Disclaimer

- Consultant for Accuray treatment planning service



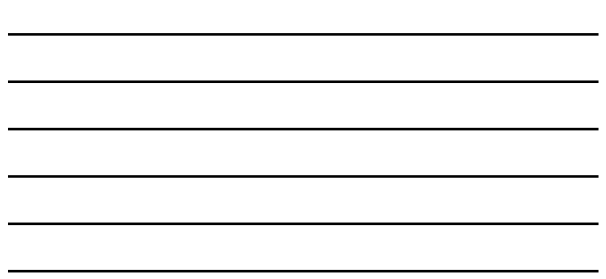
Motivation: Lung Motion

Table 2. Lung tumor motion data. The mean range of motion and the interquartile (median) range in millimeters for each subset of tumors. The number is in the parentheses (N, M, IQR).

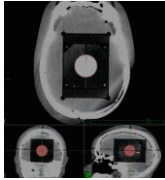
Observer	Observer		
	MI	AP	LA
Barrett ¹ Lung Les	10.0 (0-20)	--	--
Mohr ² Upper Les	7.0 (0-15)	--	--
Chen ³ L ⁴	10-30	--	--
Elting ⁵ L ⁶	7.0 (0-15)	2.0 (0-5)	2.0 (0-5)
Waggoner ⁷ L ⁸	--	--	--
Leser ⁹ Les	0-40	--	--
Mohr ¹⁰ Upper Les	0-30	--	--
Leser ¹¹ L ¹²	0-30	1.0 (0-4)	1.0 (0-1)
Waggoner ¹³ L ¹⁴	--	0	0 (0-0)
Leser ¹⁵ L ¹⁶	0-30	1.0 (0-4)	1.0 (0-1)
Waggoner ¹⁷ L ¹⁸	0-30	0 (0-0)	0 (0-0)
Barrett ¹⁹ L ²⁰	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ²¹ L ²²	8.0 (0-16)	4.0 (0-8)	1.0 (0-2)
Mohr ²³ L ²⁴	7.0 (0-14)	4.0 (0-8)	1.0 (0-2)
Waggoner ²⁵ L ²⁶	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ²⁷ L ²⁸	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ²⁹ L ³⁰	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ³¹ L ³²	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ³³ L ³⁴	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ³⁵ L ³⁶	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ³⁷ L ³⁸	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ³⁹ L ⁴⁰	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁴¹ L ⁴²	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁴³ L ⁴⁴	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁴⁵ L ⁴⁶	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁴⁷ L ⁴⁸	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁴⁹ L ⁵⁰	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁵¹ L ⁵²	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁵³ L ⁵⁴	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁵⁵ L ⁵⁶	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁵⁷ L ⁵⁸	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁵⁹ L ⁶⁰	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁶¹ L ⁶²	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁶³ L ⁶⁴	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁶⁵ L ⁶⁶	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁶⁷ L ⁶⁸	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁶⁹ L ⁷⁰	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁷¹ L ⁷²	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁷³ L ⁷⁴	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁷⁵ L ⁷⁶	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁷⁷ L ⁷⁸	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁷⁹ L ⁸⁰	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁸¹ L ⁸²	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁸³ L ⁸⁴	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁸⁵ L ⁸⁶	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁸⁷ L ⁸⁸	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁸⁹ L ⁹⁰	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁹¹ L ⁹²	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁹³ L ⁹⁴	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁹⁵ L ⁹⁶	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁹⁷ L ⁹⁸	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)
Waggoner ⁹⁹ L ¹⁰⁰	10.0 (0-20)	5.0 (0-10)	1.0 (0-1)

Table 3. Abdominal motion data. The mean range of motion and the interquartile (median) range in millimeters for each subset of tumors. The number is in the parentheses (N, M, IQR).

Site	Observer	Breathing mode	
		Normal	Deep
Pancreas	Barrett ¹ L ²	10.0 (0-20)	10.0 (0-20)
	Waggoner ³ L ⁴	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁵ L ⁶	10.0 (0-20)	10.0 (0-20)
Liver	Waggoner ⁷ L ⁸	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁹ L ¹⁰	10.0 (0-20)	10.0 (0-20)
	Waggoner ¹¹ L ¹²	10.0 (0-20)	10.0 (0-20)
Kidney	Waggoner ¹³ L ¹⁴	10.0 (0-20)	10.0 (0-20)
	Waggoner ¹⁵ L ¹⁶	10.0 (0-20)	10.0 (0-20)
	Waggoner ¹⁷ L ¹⁸	10.0 (0-20)	10.0 (0-20)
Diaphragm	Waggoner ¹⁹ L ²⁰	10.0 (0-20)	10.0 (0-20)
	Waggoner ²¹ L ²²	10.0 (0-20)	10.0 (0-20)
	Waggoner ²³ L ²⁴	10.0 (0-20)	10.0 (0-20)
Spleen	Waggoner ²⁵ L ²⁶	10.0 (0-20)	10.0 (0-20)
	Waggoner ²⁷ L ²⁸	10.0 (0-20)	10.0 (0-20)
	Waggoner ²⁹ L ³⁰	10.0 (0-20)	10.0 (0-20)
Stomach	Waggoner ³¹ L ³²	10.0 (0-20)	10.0 (0-20)
	Waggoner ³³ L ³⁴	10.0 (0-20)	10.0 (0-20)
	Waggoner ³⁵ L ³⁶	10.0 (0-20)	10.0 (0-20)
Bladder	Waggoner ³⁷ L ³⁸	10.0 (0-20)	10.0 (0-20)
	Waggoner ³⁹ L ⁴⁰	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁴¹ L ⁴²	10.0 (0-20)	10.0 (0-20)
Rectum	Waggoner ⁴³ L ⁴⁴	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁴⁵ L ⁴⁶	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁴⁷ L ⁴⁸	10.0 (0-20)	10.0 (0-20)
Vagina	Waggoner ⁴⁹ L ⁵⁰	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁵¹ L ⁵²	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁵³ L ⁵⁴	10.0 (0-20)	10.0 (0-20)
Uterus	Waggoner ⁵⁵ L ⁵⁶	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁵⁷ L ⁵⁸	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁵⁹ L ⁶⁰	10.0 (0-20)	10.0 (0-20)
Ovary	Waggoner ⁶¹ L ⁶²	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁶³ L ⁶⁴	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁶⁵ L ⁶⁶	10.0 (0-20)	10.0 (0-20)
Cervix	Waggoner ⁶⁷ L ⁶⁸	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁶⁹ L ⁷⁰	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁷¹ L ⁷²	10.0 (0-20)	10.0 (0-20)
Vulva	Waggoner ⁷³ L ⁷⁴	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁷⁵ L ⁷⁶	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁷⁷ L ⁷⁸	10.0 (0-20)	10.0 (0-20)
Penis	Waggoner ⁷⁹ L ⁸⁰	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁸¹ L ⁸²	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁸³ L ⁸⁴	10.0 (0-20)	10.0 (0-20)
Testis	Waggoner ⁸⁵ L ⁸⁶	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁸⁷ L ⁸⁸	10.0 (0-20)	10.0 (0-20)
	Waggoner ⁸⁹ L ⁹⁰	10.0 (0-20)	10.0 (0-20)

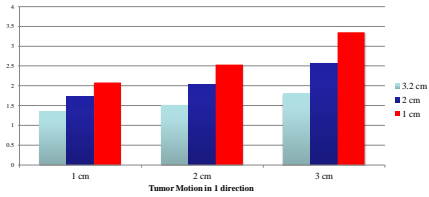


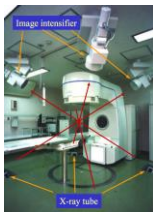
Tx Volume Comparison: 3.2 cm Spherical GTV



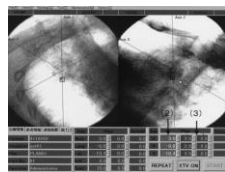
Tracking: PTV = GTV + 5mm
Motion Encompassing: PTV = ITV/iGTV + 5mm

PTV Ratio: no Tracking/with Tracking





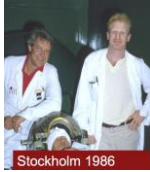
<http://rad.med.hokudai.ac.jp/>



Shirato et al. UROBP, 2000

CyberKnife System

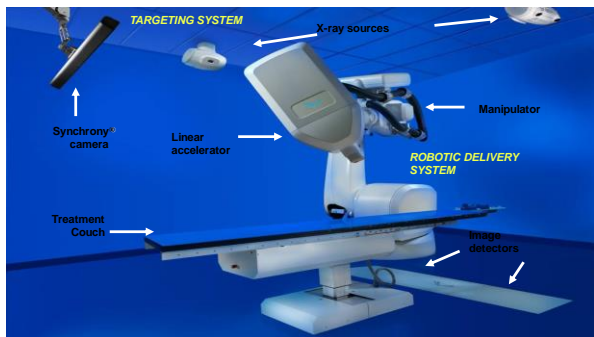
- Invented by John Adler, a neurosurgeon at Stanford University
- Had resident training with Lars Leksell in Sweden
- Noticed the GammaKnife
 - Invasive
 - Cannot fractionate
 - Only treat tumor in brain



CyberKnife System

- First CyberKnife treatment on June 4, 1994
- More than 10,000 patients treated and 244 systems installed worldwide





MD Anderson Cancer Center Cooper CyberKnife System



AAJPM Meeting 2016 10

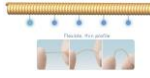
MD Anderson Cancer Center Cooper Fiducial Placement

- 4-6 fiducials are recommended
 - 1 fiducial can only track translation and at least 3 fiducial to track rotation
- Minimum 2 cm spacing between fiducials
- The triangle formed by 3 fiducials should have the smallest angle > 15 degree
- As close to the target as possible
- Ensure visibility in the 20 cm by 20 cm detector panel


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MD Anderson Cancer Center Cooper Types of Fiducial

- Gold fiducials most commonly used
 - Diameter: 0.7 to 1.2 mm
 - Length 3 to 6 mm
- Other types of fiducials offer some unique features



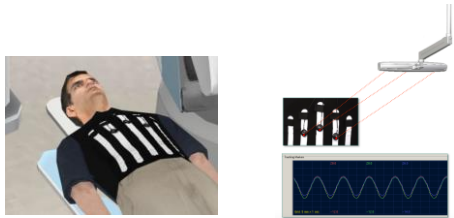
IBA Visicoil



Civco coupled markers

AAJPM Meeting 2016 12

MD Anderson Cancer Center Cooper Synchrony Vest and Camera

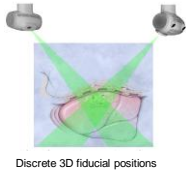


AAPM Meeting 2016

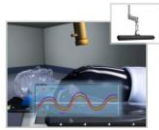
13

MD Anderson Cancer Center Cooper CyberKnife System

- How to derive real-time 3D tumor (fiducial) positions?



Discrete 3D fiducial positions



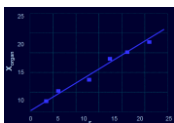
Real-time LED positions

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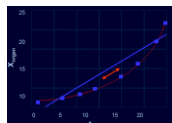
14

MD Anderson Cancer Center Cooper CyberKnife System

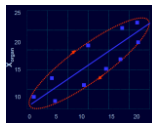
- Correlation Model – training period



Linear



Curvilinear



Dual-curvilinear

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15



Int. J. Radiation Oncology Biol. Phys., Vol. 53, No. 4, pp. 822-834, 2002
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PII S0360-3016(02)02803-1

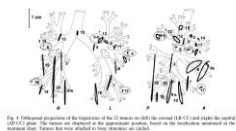
CLINICAL INVESTIGATION

Lung

PRECISE AND REAL-TIME MEASUREMENT OF 3D TUMOR MOTION IN LUNG DUE TO BREATHING AND HEARTBEAT, MEASURED DURING RADIOTHERAPY

YVETTE SEPPENWOLDE, M.Sc.,^{*} HIROKI SHIRATO, M.D., Ph.D.,[†] KEI KITAMURA, M.D., Ph.D.,[†] SHINICHI SHIMIZU, M.D., Ph.D.,[†] MARCEL VAN HERK, Ph.D.,^{*} JOOS V. LEBESQUE, M.D., Ph.D.,^{*} AND KAZUO MIYASAKA, M.D., Ph.D.[†]

^{*}Department of Radiotherapy, The Netherlands Cancer Institute, Antoni van Leeuwenhoek Hospital, Amsterdam, The Netherlands; [†]Department of Radiation Medicine, Hokkaido University School of Medicine, Sapporo, Japan



- 20 patients were recruited from Mitsubishi/Hokkaido RTRT treatment
- Hysteresis effect was observed in 10 of 21 tumors (1-5 mm)
 - "The tumor followed a different path during inhalation than during exhalation."
- Mostly in the sagittal plane

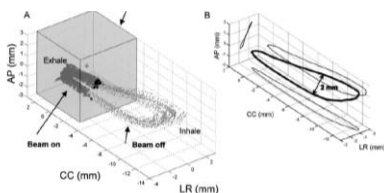


Fig. 3. (A) The 3D path of the tumor of Patient 10 during one treatment portal. The gray dots represent the tumor position throughout the treatment, and the black dots represent the tumor position as the tumor is detected to be inside the RTRT range (transparent box). The asterisk represents the planned zero position. (B) The average 3D curve of the



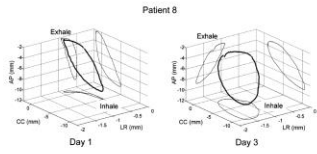


Fig. 5. The 3D trajectory of the tumor of Patient 8: (A) On Day 1, the pattern was similar to that on Day 2 and 4. (B) Day 3. The shape and direction of the motion changed between subsequent days. In the coronal plane on Day 3, hysteresis was present, whereas on Days 1, 2, and 4, this was not the case. In the transaxial (AP-LR) plane, the motion switched from left to right on Days 1, 2, and 4 and to right to left on Day 3.

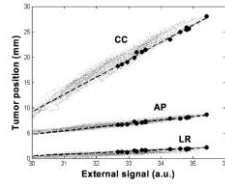


• Correlation Model – linear

(1) A linear model, that is simply a linear fit between target positions (3D) and scalar external marker positions (t),

$$\begin{pmatrix} x \\ y \\ z \end{pmatrix}_{\text{Target}} = \begin{pmatrix} A_x \\ A_y \\ A_z \end{pmatrix} r + \begin{pmatrix} B_x \\ B_y \\ B_z \end{pmatrix}$$

Coefficients (A, B) are found by least-squares fitting to image/marker data-points (Fig. 3).



Seppenwoolde et al. Med. Phys., 2007

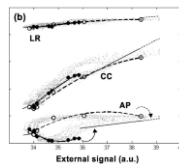


• Correlation Model – Dual-curvilinear

(2) A polynomial model with two fitted second-order polynomials, one through the image/marker data-points that are determined to be in the inhale phase of the breathing cycle, and one that is determined for the exhale phase.

$$\begin{pmatrix} x \\ y \\ z \end{pmatrix}_{\text{Target}} = \begin{pmatrix} A_x \\ A_y \\ A_z \end{pmatrix} r^2 + \begin{pmatrix} B_x \\ B_y \\ B_z \end{pmatrix} r + \begin{pmatrix} C_x \\ C_y \\ C_z \end{pmatrix}$$

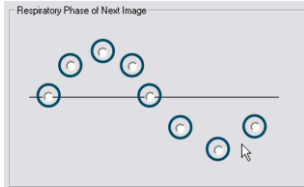
An example is shown in Fig. 4.



Seppenwoolde et al. Med. Phys., 2007



- It's recommended acquisition of model points covering whole breathing period.

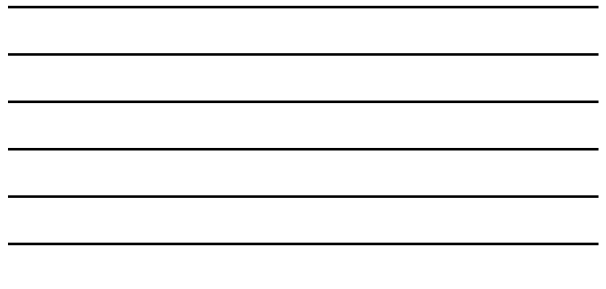
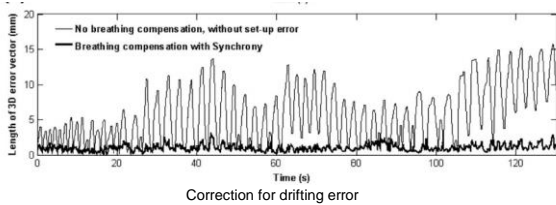


- Multiple breathing models are allowed
 - Abrupt breathing pattern change
 - Irregular breathing pattern, e.g. cough
 - Patient moves
 - Patient wants a break
 - Different direction could have different model

- For the same model during treatment, the model points keep updated once a pair of x-ray images is taken - adaptive
 - To account for slow change of breathing pattern or drifting
 - The oldest model point will be discarded first – last in last out
- Prediction is used to account for system latency (115 ms)
 - Autoaggressive process – normalized least mean square (nLMS)
 - The predicted tumor position is compared to the real detected position
 - Once the difference is beyond a threshold, an E-stop occurs

- CyberKnife doesn't take continuous x-ray image pairs so the ground truth between pairs is missing
- Data from Mitsubishi/NTT hospital RTRT system used to retrospectively evaluate CyberKnife modeling accuracy
 - 8 lung cancer patients
 - Peak to peak motion > 1 cm
 - Synchronized recordings of internal tumor/fiducials motion and external markers motion
- All three models were evaluated to find the optimal model
 - Linear, polynomial and hybrid models

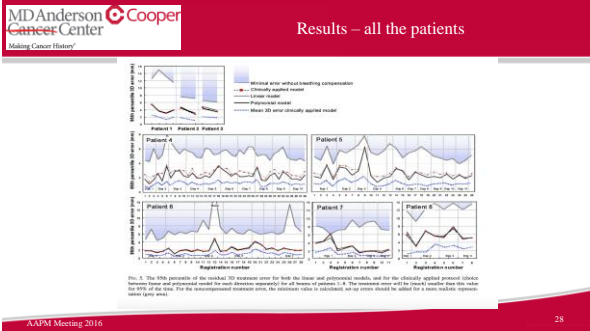
Seppenwoolde et al. Med. Phys., 2007

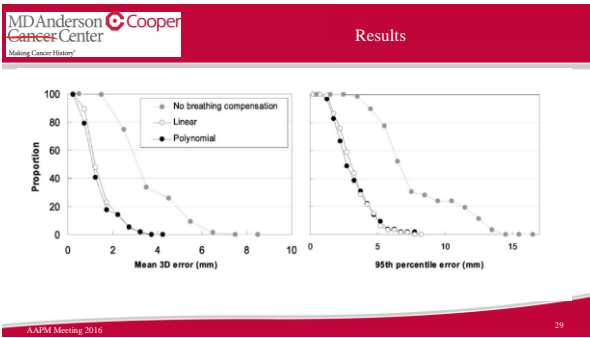


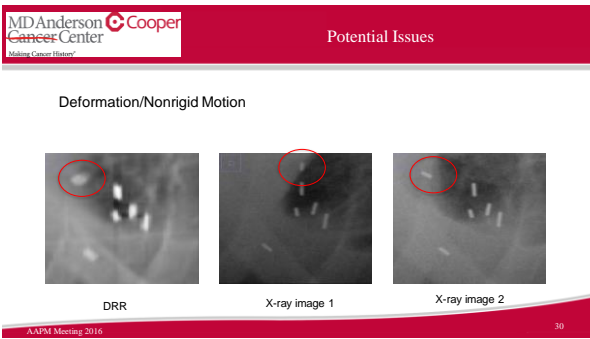
Patient 1	Patient 2	Patient 3	Patient 4
LR	CC	AP	LR
CC	AP	LR	CC
AP	LR	CC	AP
LR	CC	AP	LR
CC	AP	LR	CC
AP	LR	CC	AP
LR	CC	AP	LR
CC	AP	LR	CC
AP	LR	CC	AP

Fig. 8. Examples of the range of internal and external motion errors (1SD), arranged by each patient, for the reconstructed error and for the polynomial regression error (polynomial regression error) for different set-up errors in the external motion position (LR, CC, AP) for the 10 patients.

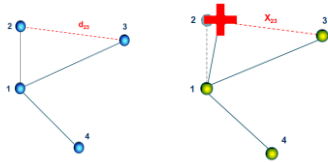








Deformation/Nonrigid Motion – Rigid Body Error: d23-x23



- System is capable to correct 1.5° in roll, 1.5° in pitch and 3.0° in yaw
- 25 patients, 2796 pairs of images were analyzed
- PTV were created by adding 3 mm (UL) and 5 mm (LL) margins
- CTV were rotated based on reported angles + random shifts
- For 94.4% and 97.1% of the the images in LL and UL, the CTV is 100% covered by the PTV and rest mean CTV coverage were 95.6% (LL) and 99% (UL)

	Roll	Pitch	Yaw
Lower lobe	0.25° ± 5.7°	-0.21° ± 7.3°	0.23° ± 5.3°
Upper lobe	0.40° ± 2.1°	0.05° ± 1.8°	0.1° ± 2.1°

Xu et al. AAPM 2013



Summary

- Multiple systems with RTRT capability were reviewed
- Three breathing models were introduced
- Prediction uncertainties were analyzed
- Real-time tumor tracking with fiducials significantly reduce PTV and nearby healthy tissue irradiated
- Deformation and rotation could introduce uncertainties

Thank you!