





5 NRG Oncology Specific Aims

Conduct practice-defining research for the major **gender-specific malignancies** (breast & gynecologic cancers & prostate cancer) while capitalizing on common biologic features and interactive research opportunities among these diseases;



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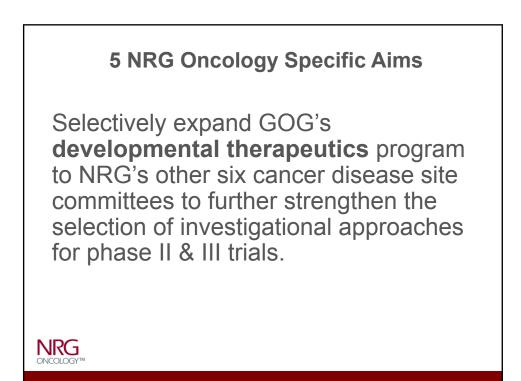
Investigate new developments in **medical technology**, including radiation oncology, imaging, surgical technology, & IT, for opportunities to advance the care of patients with localized / locally advanced cancers;

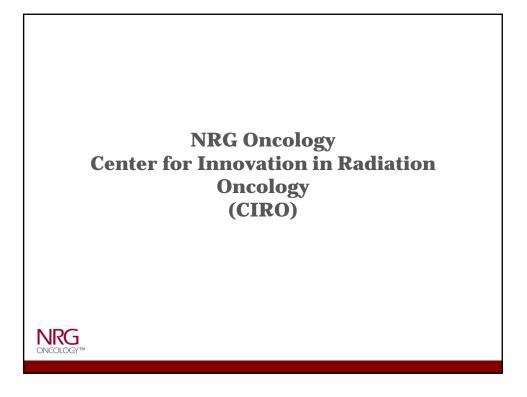


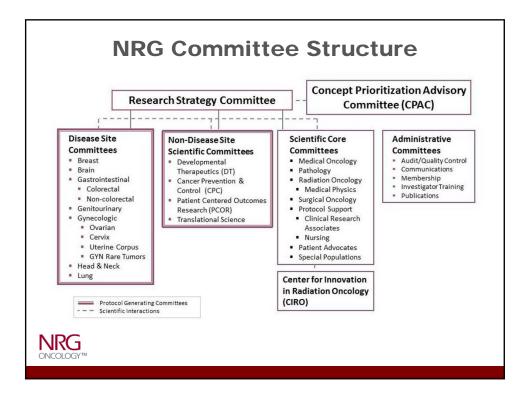
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Integrate and expand the legacy groups' translational science programs to better inform biomarker- and biologic pathway-defined approaches to risk stratification, investigational therapy assignment, & clinical trial decisionmaking;





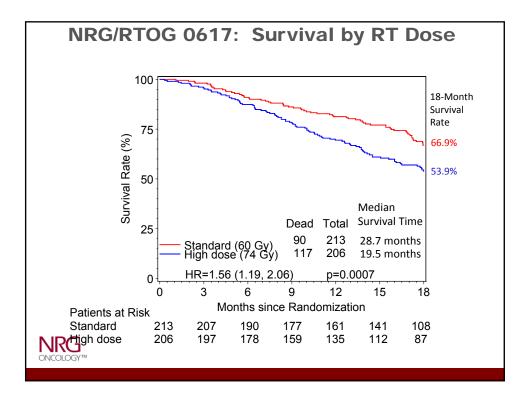


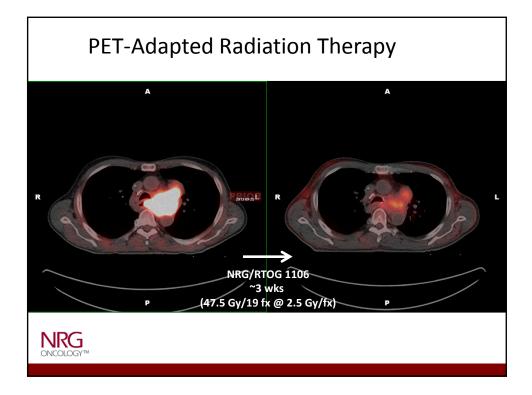


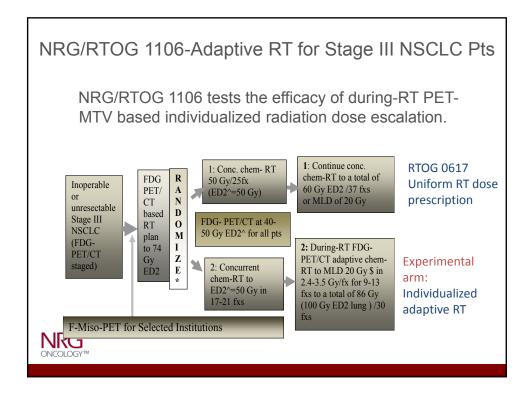
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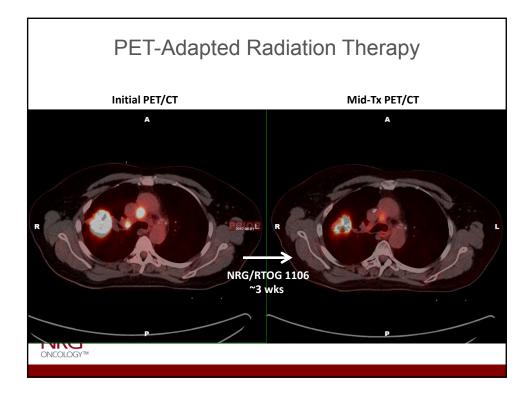
- Promote innovative RT research within all NCTN
- Accelerate testing new rad onc innovations in NCTN
- · Facilitate innovation in all appropriate protocols
- Foster intergroup & protocol harmonization
- · Reduce timelines for development of new protocols
- · Improve the clarity of NCTN protocols

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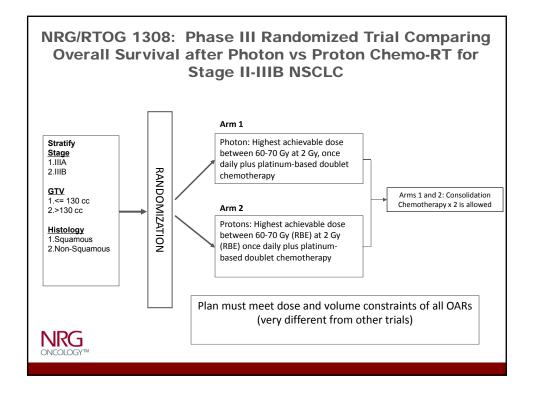


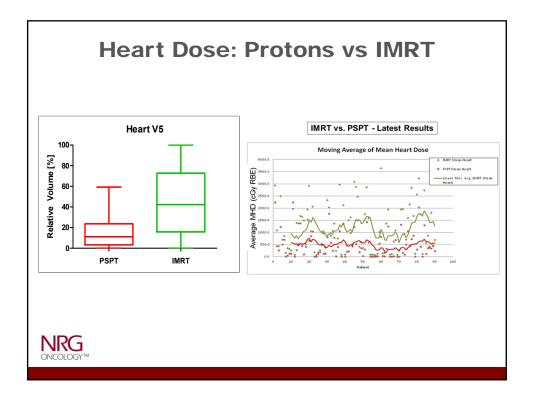


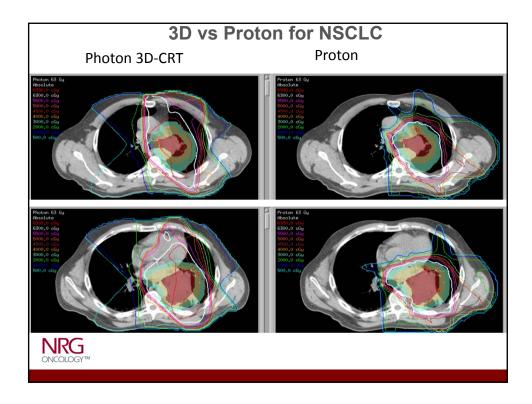


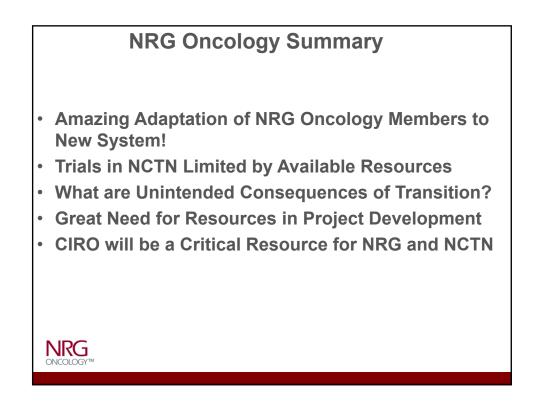












SBRT Protocols-Past&Present Examples



Stereotactic Body Radiation Therapy for Inoperable Early Stage

Lung Cancer

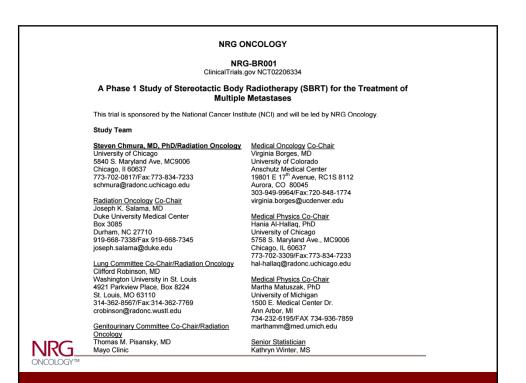
Robert Timmerman, M.D., Rebecca Paulus, B.S., James Galvin, Ph.D., Jeffrey Michalski, M.D., William Straube, Ph.D., Jeffrey Bradley, M.D., Achilles Fakiris, M.D., Andrea Bezjak, M.D., Gregory Videtic, M.D., David Johnstone, M.D., Jack Fowler, Ph.D., Elizabeth Gore, M.D., and Hak Choy, M.D.

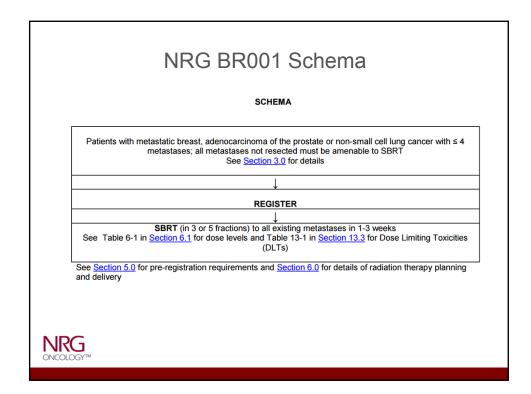
Abstract

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Context—Patients with early stage but medically inoperable lung cancer patients have a poor rate of primary tumor control (30-40%) and a high rate of mortality (3-year survival 20-35%) with current management.

Objective—To evaluate the toxicity and efficacy of stereotactic body radiation therapy in a high risk population of patients with early stage but medically inoperable lung cancer.





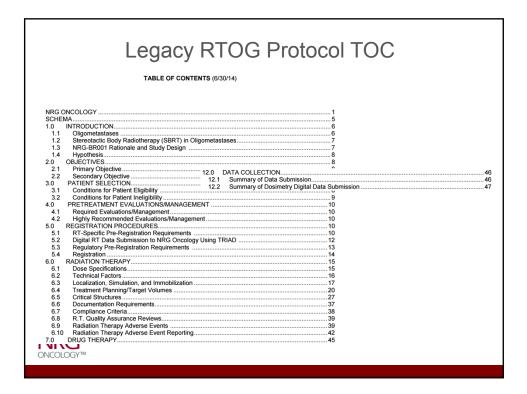


	Table Of C	ontent		
			-	
SCHEMA			RATION, STUDY ENTRY, AND WITHDRAWAL PROCEDURES	
1. OBJECTI	VES	8.1 II 8.2	vestigator Registration Requirements	
	rimary Objective	8.2	Site Registration Requirements RT-Specific Pre-registration Requirements	
	econdary Objectives	8.4	Patient Enrollment	
2. BACKGF	OUND	DDU	G INFORMATION	
2.1	Rationale for Arm 1: Modestly 9.	9.1	Investigational Study Agent	
	Chemotherapy (60 Gy/30 fract	9.1	Commercial Agent: Cisplatin	
	40/mg/m ²)			
2.2	Rationale for Arm 1 Radiosens 10.1	BIOMA	RKER, CORRELATIVE, AND SPECIAL STUDIES	
	mg/m	10.1	Biomarkers	,
2.3	Rationale for Arm 2: Modestly	10.2	Ouality of Life	
	Therapy (60 Gy/30 fractions/5	10.3	FDG-PET/CT Imaging	
2.4	Measuring Patient-Reported O		00	
2.5	FDG-PET/CT Imaging as a Pre 11.	MOD	ALITY REVIEWS	
	Control and Progression-Free §	11.1	Radiation Therapy Quality Assurance Reviews	
2.6	Circulating HPV DNA as a Pot	11.2	Drug Quality Assurance Reviews	8
2.7	PI3K Pathway Activation and 1			
	Prognostic Biomarkers of HPV 12.		A AND RECORDS	
		12.1	Data Management/Collection	
	T SELECTION, ELIGIBILITY, #	12.2	Summary of Data Submission	
3.1	Patient Selection Guidelines	12.3	Digital Data Submission Requirements	
3.2 3.3	Eligibility Criteria	12.4		
4. REQ	UIREMENTS FOR STUDY ENTRY,	TREAT	MENT, AND FOLLOW-UP35	
5. TREATM	IENT PLAN/REGIMEN DESCRIPTIO			
5.1				
5.2				
5.3	Surgerv			



appropriate te	chnology for this protocol can be directed	are given in Table 5.2.1A. Questions regardin I to the protocol PI or medical physics co-chair
	<u>Summary</u> of Treatment Technology I	
Technology Beam Modality	Requirement MV Photons	Comments Cobalt-60 & Linac Allowed; Charged particle beams (including electrons, protons, and heavier ions) are not allowed
Beam Energy	1 to 18 MV	Minimize use of high energy in lung. 6 MV or lower energies should be predominately used in low-density tissue.
Treatment Technique	3DCRT (static, arc) or intensity modulated techniques (IMRT, VMAT)	Tomographic and robotic techniques allowed
Image Guidance	Treatment Machine must be equipped to provide daily image guidance. The minimum required image guidance techniques as a function of treatment techniques are given in Section 5.2.11.	Non-ionizing guidance is allowed, but secondary image verification is required

•	mmobilization and Simulation
Topic/Parameter	y of General Simulation Guidelines Guideline
Immobilization	Proper immobilization with appropriate clinical devices to ensure reproducibility is required. Patient discomfort should be minimized.
Motion Control	Recommended in cases where the extent of motion and patient geometry may result in a violation of <u>dosimetric</u> constraints to normal tissues. Institutions should use their clinical judgment when determining which patients are appropriate for a motion management technique.
CT Slice Thickness	2 mm or less is recommended. No more than 3 mm shall be used in the vicinity of the target. PTV size should be taken into consideration when choosing the slice thickness. Slices of 1-2 mm should be used for tumors that are 1 cm or less in the largest dimension.
Use of Contrast	IV and/or oral contrast can be used at the clinical discretion of the treating institution. It is recommended to perform a non-contrast enhanced CT for planning. For how to handle treatment planning on a contrast CT, please see the treatment planning section.

To simplify the simulation and planning process, Table 5.2.2B highlights the recommended and minimum requirement for motion assessment, treatment planning imaging, and PTV margins for all SBRT protocols.



Treatment Technique	Recommended Method for Motion Assessment During Simulation	Minimum Method for Motion Assessment During Simulation	Scan(s) Required for Treatment Planning	Additional Instructions
Free breathing treatment using an ITV approach, including abdominal compression	4DCT	CT scans at normal inhale and exhale positions	Average/Untagged scan from 4DCT for dose calculations; MIP may be desirable to aid ITV definition; If 4DCT not available, planning scan should be at normal exhale	A free breathing non-4DCT scan is not appropriate
Gating with a gating window	4DCT	Exhale CT plus fluoro (free-breathing + fluoro strongly discouraged due to baseline shift)	Reconstructed average of gating window scans if 4DCT or normal exhale scan	Exhale recommended since it gives the most conservative measure of lung dosimetry
Gating with breath hold (i.e. ABC)	Reproducibility of breath hold confirmed (examples: multiple low dose scans over tumor, repeat fluoroscopy or scout images)	N/A	Scan in breath hold position	Inhale recommended since it maximizes lung volume
Tracking	4DCT or breath hold CT	N/A	4DCT or breath hold CT	Need to know tumor trajectory

	Plan	Mean (Gy)	Standard error (Gy)	Comparison	P value	plans, MII	p lans	, and AIP pla	ns.		
Dmax	FB	54.4	8.6	FB vs MIP	0.116		Plan	Mean (cm ³)	Standard error (cm3)	Comparison	P valu
Dmax	MIP	54.8	8.7	MIP vs AIP	0.132	Abs, V5	FB	603.9	336.0	FB vs MIP	< 0.00
	AIP	54.5	8.5	FB vs AIP	0.522	AUS. V.J	МР	550.7	320.5	MIP vs MIP	< 0.00
D _{min}	FB	44.0	6.3	FB vs MIP	< 0.001		AIP	603.0	337.5	FB vs AIP	< 0.00
- min	MIP	45.0	6.5	MIP vs AIP	0.006	Abs. V10		323.7	182.9	FB vs MIP	<0.00
	AIP	44.4	6.3	FB vs AIP	0.003	Abs. VIU	MIP	304.0	182.9	MIP vs MIP	< 0.00
Dmean	FB	50.3	7.1	FB vs MIP	0.002		AIP	324.5	186.5	FB vs AIP	<0.0
20 mean	MIP	50.9	7.3	MIP vs AIP	0.008	Abs. V20		167.7	117.3	FB vs MIP	0.0
	AIP	50.4	7.1	FB vs AIP	0.017	A08. V20	MIP	160.1	117.5	MIP vs AIP	0.00
D95	FB	47.3	6.3	FB vs MIP	0.001		AIP	167.7	114.5	FB vs AIP	0.9
2.70	MIP	48.1	6.5	MIP vs AIP	0.006	Abs. V30	FB	91.9	73.1	FB vs MIP	0.0
	AIP	47.5	6.4	FB vs AIP	0.003	A08. V 50	MIP	89.2	71.8	MIP vs AIP	0.04
D90	FB	48.0	6.4	FB vs MIP	0.001		AIP	92.0	74.2	FB vs AIP	0.9
570	MIP	48.7	6.6	MIP vs AIP	0.006	Abs, V35	FB	92.0 66.4	55.2	FB vs MIP	0.1
	AIP	48.2	6.4	FB vs AIP	0.010	A08. V 55	гь МІР	64.9	54.5	MIP vs AIP	0.1
CI	FB	0.71	0.09	FB vs MIP	0.010		AIP	66.3	55.9	FB vs AIP	0.8
0.	MIP	0.67	0.11	MIP vs AIP	0.111	Abs, V40		46.0	40.4	FB vs MIP	0.8
	AIP	0.69	0.09	FB vs AIP	0.002	Abs. V40	MIP	46.0	40.4	MIP vs AIP	0.5
TV _{PD}	FB	49.9	33.9	FB vs MIP	0.694		AIP	45.8	40.2	FB vs AIP	0.5
1.10	MIP	50.1	33.7	MIP vs AIP	0.437		AIF	43.0	40.9	FD VS AIF	0.50
	AIP	49.6	33.9	FB vs AIP	0.035						
V _{PD}	FB	66.4	42.1	FB vs MIP	0.070						
• rD	MIP	69.5	40.7	MIP vs AIP	0.154						
	AIP	67.0	41.9	FB vs AIP	0.147						

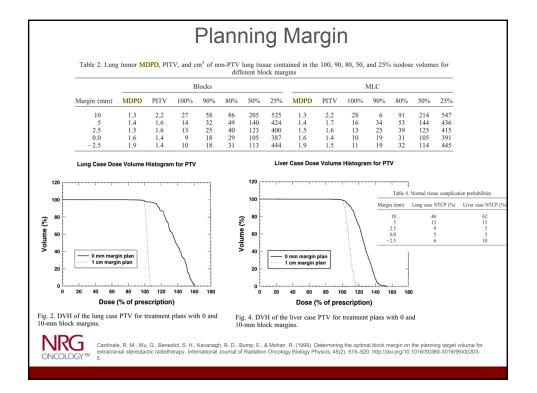
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	Description GTV to receive XXXX cGy	
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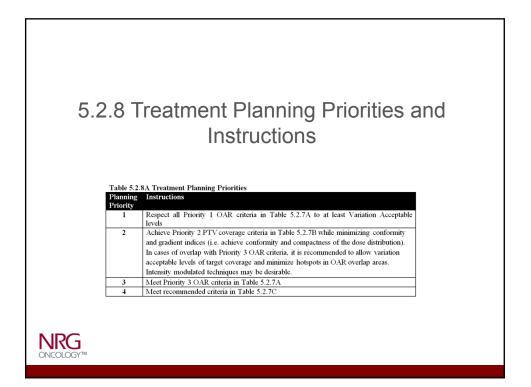
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5.2.	6Treatment Planning Guidelines
Topic/Parameter	Guidelines
Planning Technique	3DCRT, conformal arc, and intensity-modulated techniques (IMRT, VMAT) allowed. Tomographic and robotic techniques also allowed.
Number of Beams	As planning dictates although ≥7 beams are recommended for static beam plans due to skin toxicity considerations. Similarly, arcs should cover an appropriate range so as to deliver a safe dose to the skin.
Beam Arrangement	Coplanar or non-planar, non-overlapping, non-opposing beams or arc therapy (non-coplanar arcs allowed). Combination of static and arc beams allowed.
Beam Energy	As planning dictates although lower energies preferred for lung
Block Margin (for 3DCRT	0-2 mm
Minimum Field Size	As planning dictates although only the smallest field size accurately commissioned (e.g. small field output factors are within 5% of published standards or values) at the institution should be used. Because of concerns with small field dosimetry, field sizes above 2 cm x 2 cm are preferable.
Dataset for Dose Calculation	ITV Approach – Average from 4DCT or normal exhale if 4DCT not available (Free breathing CT is not appropriate) Breath Hold – CT taken at treatment breath hold Gated – Average from gating window phases from 4DCT or the median phase in the gating
	window Tracking – 4DCT or breath hold CT
	Contrast Scans are not recommended for dose calculations. Recommend obtaining a non- contrast scan during simulation for dose calculation. If a contrast scan is used for dose calculation, density/material overrides are recommended when dose calculation accuracy may be affected.
Dose Calculation Algorit	Modern algorithms that accurately handle tissue heterogeneity and scatter should be used. IROC maintains an updated list of approved algorithms. Density corrections must be applied. Density overrides of the ITV are not recommended for photon treatment. All doses should be reported in terms of dose-to-water and not in terms of dose-to-medium.
Dose Grid Resolution	2 mm x 2 mm dose grid resolution or smaller is strongly recommended.



FS (am)	N		PTV D	95 (%)			MLD	(%)	
FS (cm)	N	EPL-3D	AAA	ĊĊĊ	MC	EPL-3D	AAA	ČCC	MC
$3 \leq FS < 5$	50	95.3±1.9	82.4±5.1	82.6±6.2	82.3±6.0	94.4±7.4	90.7±5.6	91.8±6.1	90.9±5.7
$5 \leq FS < 7$	62	95.8±2.1	85.3±5.2	85.7±6.1	85.6 ± 5.8	100.5 ± 2.5	95.3±2.7	96.1±2.4	96.1±1.8
7≤FS<10	21	95.9±1.7	90.4±3.7	90.6±3.8	90.8±3.8	102.0 ± 2.8	96.1±2.1	95.0±2.4	97.3±3.2
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isla Location	,		PTV	/ D95 (%)		EPL-3D		LD (%)	МС
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	N d 39	EPL-3	PTV D AAA .0 81.6±4	D95 (%) CCC .4 81.4±5.	MC .8 81.4±5.	EPL-3D 8 97.2±6.1	AAA 92.1±5.	LD (%) CCC	5 92.9±5.3
Location Lung-islan	N d 39 l 44	EPL-3 9 95.2±2. 4 96.5±1.	PTV D AAA .0 81.6±4 .8 86.8±4	D95 (%) CCC .4 81.4±5. .9 87.4±5.	MC .8 81.4±5.1 .6 86.9±5.1	EPL-3D 8 97.2±6.1	AAA 1 92.1±5. 7 94.3±4.	LD (%) CCC 1 92.9±5.6 8 94.4±4.3	5 92.9±5.3 94.9±4.9

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Name of	Dosimetric	Priorit		Variation	Notes (if		
Structure	parameter*		Protoco	ol Acceptable	needed)		
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Table 5.2.7E	Normal Structur	e Constraints an	d Complianc	e Criteria <mark>(**T</mark>	HESE ARE JUST		
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	D'mean(Gy)	+				-	
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Table 5.2.70	Additional Targe	et and Normal S	tructure Reco	mmendations –	Not to be used in		
					l be determined by		
the protocol					<mark>l when possible**</mark>)		
	Dosimetric	Recomme	andation Table 5.2.7D	Notes Delivery Compli	anco Critorio		
Name of	parameter		Delivery Metr		Per Protocol	Variation	Notes
Structure		2000			1 01 1 1000001	Acceptable	1.0000
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Structure	$\frac{D_{mean}(Gy)}{D_{1cc} (Gy)}$	<u>≤105%</u>	Start date (X d	ays/weeks after 2			
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5.2.9 Patient specific QA

Any patient-specific QA that needs to be acquired should follow institutional guidelines. For intensity modulated techniques, patient specific QA is highly recommended.

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Treatment Technique	Acceptable Methods for Daily Image Guidance	Matching Instructions
ITV/free breathing (includes abdominal compression)	Volumetric Imaging Volumetric Imaging (i.e. CBCT or CT on rails) is strongly recommended	 Initial rigid alignment followed by soft tissue match with average CT and slow CBCT 4DCT to 4D-CBCT can be used when capability exists
	Planar Imaging If volumetric imaging is not available, then an appropriate tumor surrogate (i.e. implanted fiducials) must be able to be accurately imaged in the treatment position with 2D imaging. The patient surface is not an appropriate surrogate for tumor setup although surface based imaging may be used during treatment to assess unexpected patient motion. Note that when orthogonal 2D imaging (with or without implanted fiducials) is employed for sites where respiratory motion is expected and not controlled via motion management techniques, care must be taken to ensure accurate tarceting of the ITV within the treatment. For example, static kV imaging at an	Rigid alignment to bony anatomy Repeat imaging to ensure tumor surrogate is within ITV Repeat imaging at each treatment port to ensure tumor surrogate remains within the ITV is very strongly recommended
	angening on the FTV within the treatment. FO example, statuc kV maging at an undetermined breath hold position would not be adequate IGRT for treating a free-breathing lung turnor. Repeat imaging during treatment is recommended to verify that the turnor is in the ITV If any significant baseline shifts are noted, resimulation should be strongly considered	
Gating with a gating window	The baseline gating position/phase should be verified using appropriate imaging techniques Volumetric Imaging (i.e. CBCT or CT on rails) is strongly recommended for the initial localization to verify isocenter and tumor trajectory	 Initial rigid alignment followed by soft tissue match for baseline gating position
Gating with breath hold (ie ABC)	Volumetric imaging recommended; planar at breath hold position acceptable – repeated imaging recommended to ensure reproducibility of breath hold All imaging should be done at breath hold treatment position	 Initial rigid alignment followed by soft tissue match of tumor or surrogate
Tracking	Volumetric imaging or real-time fluoroscopic imaging of tumor surrogate required based on treatment machine capabilities.	 Initial rigid alignment followed by soft tissue match of tumor or surrogate in baseline position

