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Table S1. Anatomical boundaries of chest wall and whole breast CTV

	Chest wall	Whole breast
Cranial	Guided by palpable /visible signs; if appropriate guided by the contralateral breast; maximally up to the inferior edge of the sternoclavicular joint	Upper border of palpable /visible breast tissue; maximally up to the inferior edge of the sternoclavicular joint
Caudal	Guided by palpable/visible signs; if appropriate guide by the contralateral breast	Clinical reference; most caudal CT slice with visible breast
Anterior	Skin	5mm under skin surface
Posterior	Include pectoralis muscles, chest wall muscles, excluding ribs	Exclude pectoralis muscles, chest wall muscles, ribs
Lateral	Guided by palpable/visible signs; if appropriate guide by the contralateral breast. Usually anterior to the mid-axillary line, excludes lattices dorsi muscle	Lateral breast fold; anterior to the lateral thoracic artery
Medial	Lateral to the medial perforating thoracic internal vessels; maximally to the edge of the sternum	Lateral to the medial perforating thoracic internal vessels; maximally to the edge of the sternum

Notes: For patients with stage III according to AJCC 8th edition, the posterior of the whole breast can extend to the rib interface.

Table S2. Anatomical boundaries of regional nodal CTV

	Cranial	Caudal	Anterior	Posterior	Lateral	Medial
Medial Supra-clavicular	Caudal to the cricoid cartilage	Junction of brachioceph-axillary veins / Caudal edge clavicle head	Posterior edge of sternocleidomastoid muscle	Posterior surface of internal carotid artery or anterior surface of scalene muscles	Medial border of clavicle head or 5 mm below the skin surface	Medial border of internal carotid vessels
Lateral Supra-clavicular*	Caudal to the cricoid cartilage	Junction of Brachiocephaxillary veins. /Caudal edge clavicle head	Posterior surface of internal carotid artery or anterior surface of scalene muscles	At the trapezius	Medial border of clavicle head or 5mm below the skin surface	At the longus coli
Internal mammary	Junction of Brachioceph-axillary veins.	Cranial side of the 4rd rib (in selected cases 5th rib)	Ventral limit of the vascular area	Pleura	7mm from internal mammary vessels	7mm from internal mammary vessels
Axilla-level I	Below the humeral head	At the point where the pectoralis major inserts on the ribs (around 4th to 5th rib)	At posterior pectoralis major & minor muscles	An imaginary line anterior to the surface of latissimus dorsi muscle, subscapularis muscle and teres major muscle	Cranially up to an imaginary line between the major pectoral and deltoid muscles, and further caudal up to a line between the major	Lateral border of pectoralis minor musle

					pectoral and latissimus dorsi muscles	
Axilla-level II	Insertion point of pectoralis minor muscle	The caudal border of the pectoralis minor muscle	Pectoralis minor muscle	Anterior border of ribs and serratus anterior	Lateral edge of pectoralis minor muscle	Medial edge of pectoralis minor muscle
Axilla-level III	Pectoralis minor muscle insert on Coracoid	Axillary vessels cross medial edge of pectoralis minor muscle	Posterior surface of pectoralis major muscle	Anterior border of ribs or posterior border of subclavian or axillary vessels	Medial border of pectoralis minor muscle	Lateral border of clavicle, ribs, junction of brachiocephaxillary veins.

Notes:

- 1) The regional nodal CTV target volume should include the medial supra-clavicular, axilla-level III, undissected axilla-level II, and internal mammary lymph nodes.
- 2) The inclusion of axilla-level I and II in the regional nodal CTV is at the discretion of radiation oncologists based on the clinical treatment indications. The rotter's nodes are included in the CTV target volume of axilla-level II.
- 3) For N2, N3, or high-risk N1 patients, it is recommended to include the lateral supra-clavicular nodes in the regional nodal CTV
- 4) When the tumor is in the inner lower quadrant, the caudal border of internal mammary nodes is recommended to extend to the cranial side of the 5th rib.

Table S3. Tumor bed GTV and CTV contours

GTV	Includes seroma and/or surgical clips
CTV	GTV with a margin of 1 cm in all directions and without exceeding CTV of the whole breast

Abbreviations: GTV=gross tumor volume, CTV=clinical target volume

Table S4. Dosimetric coverage for target volume

Target volume		Requirements	
	Per Protocol	D95%>26Gy	
	Acceptable variation	D90%>26Gy	
PTV of chest wall/whole breast and	Per Protocol	V28Gy <5%	
RNI	Acceptable variation	V29Gy <5%	
	Per Protocol	V25Gy >99%	
	Acceptable variation	V24Gy >99%	
CI for PTV of chest wall/whole	Per Protocol	≥0.95 , ≤2	
breast and RNI	Acceptable variation	≥0.85 , ≤2.5	
	Per Protocol	D95%>36.4Gy	
	Acceptable variation	D90%>36.4Gy	
Tumor bed PTV	Per Protocol	V39Gy <5%	
	Acceptable variation	V40Gy <5%	
	Per Protocol	V35Gy >99%	

	Acceptable variation	V34Gy >99%	
	Per Protocol	≥0.95 , ≤2.5	
CI for Tumor bed PTV	Acceptable variation	≥0.9 , ≤3	

Notes:

- 1) When evaluating the treatment plan of chest wall/whole breast with RNI, tumor bed boost doses should not be considered.
- 2) CI=Conformity index, the volume covered by 95% of the prescription dose / PTV volume
- 3) For evaluation of IMRT plan: a) The whole breast CTV is expanded by 5-8mm in three dimensions to form the PTV. The PTV is limited to 5mm below the skin and the surface of the pectoral muscle or chest wall to form the whole breast PTV_eval for IMRT plan evaluation; 2) The chest wall and regional nodal CTV is expanded by 5-8mm in three dimensions to form an integrated PTV; Limit the PTV to the skin as the anterior boundary and the chest pleura as the posterior boundary to form the PTV_eval for plan evaluation.
- 4) For evaluation of proton plan: a) the dosimetric coverage of the CTV is evaluated; b) The chest wall CTV is recommended to retract 5mm from skin; c) If the chest wall is very thin, 3mm can be considered.

Table S5. DVH constraints for OARs

CAR		The ARROW trial	
OARs	Dosimetric parameter	Per protocol	Acceptable variation
<u>-</u>	Mean	<3Gy	<3.5Gy
TI (C. 1.C. 1.1.1.	V15Gy	<10%	<15%
Heart for left-sided breast cancer	V4Gy	<20%	<25%
	N/A	N/A	N/A
T . (C . 1 . 1 . 1	Mean	<1Gy	<1.5Gy
Heart for right-sided breast cancer	V2Gy	<15%	<20%
<u>-</u>	Mean	<7.5Gy	<8Gy
	V4Gy	<45%	<50%
Ipsilateral Lung	V10Gy	<30%	<35%
	V15Gy	<20%	<23%
G I II	Mean	<1Gy	<1.5Gy
Contralateral lung	V2Gy	<10%	<15%
Spinal cord	Max	<20Gy	N/A
Ipsilateral Humeral head	Mean	<10Gy	<11Gy

Supplemental material

Contralateral breast	Mean	<3.5Gy	<5Gy
Brachial plexus	D0.1cc	<27Gy	<28Gy
Position	D1cc	< 26Gy	< 27Gy
Esophagus	D 0.01cc	< 27Gy	< 28Gy
Thyroid	Mean dose	< 10Gy	< 11Gy
Skin ring	V2Gy	<10%	<15%

Supplemental material

NCT	Coun	Age	Start	Random	Enrolled	Reconstru	Tumor bed	IMNI	Study	Control	Sample	Primary	Status
Number	try	(years)	year	ized	patients	ction	Boost		group	group	size	Endpoint	
WBI alone													
NCT0558 6256	Italy	18-99	2021	No	Indicated for WBI	N/A	Optional, SIB of 30Gy/5Fx or sequential 7.6Gy/2Fx	N/A	26Gy/5F x/1w	N/A	300	Acute and chronic toxicity	Recruit ing
NCT0531 8274	Mexi co	≥18	2021	Yes	BCS, pTis-T2, N0	N/A	Study group: No boost; Control group: SIB of 48Gy/16Fx in high risk patients	N/A	26Gy/5F x/1w	42.5Gy/16 Fx	72	Local control	Recruit
NCT0466 9873	Brazil	50-90	2021	Yes	BCS, unifocal disease, T < 3cm, pN0, negative LVSI, Grade 1-2, margin >	N/A	No	N/A	Group1: 26Gy/5F x/1w for WBI Group2: 26Gy/5F x/1w for PBI	40Gy/15F x/3w for WBI	36	Local recurrence	Recruit

					2mm								
NCT0541 7516 RNI	Cana da	≥50	2023	Yes	BSC, negative margin, N0	N/A	No	N/A	26Gy/5F x/1w for PBI	26Gy/5Fx /1w for WBI	910	Local recurrence	Recruit ing
NCT0378 8213	India	≥18	2019	Yes	BCS, or after NAC, or mastectomy with pT3/T4 or pN2 or pT0-2pN0-1 with a Cambridge Score of 3 or more	Yes	Study group:SIB of 32Gy/5Fx, or sequential 12Gy/4Fx; Control group: SIB of 48Gy/15Fx, or sequential 12Gy/4Fx	Based on instituti onal policy	26Gy/5F x/1w for WBI/Che st wall, and RNI in N+ or after NAC	40Gy/15F x/3w for WBI/Ches t wall, and RNI in N+ or after NAC	2100	Locoregional recurrence rate	Recruit
NCT0464 8904	Unite d States	≥30	2020	No	mastectomy with reconstructi on; pathologic T0N1a-2a, T1N1a-2a, T2N1a-2a, T3N0-2a, all	Yes	Optional chest wall boost of 5.2 Gy for 1-2 fractions or 2.5 Gy for 1-4 fractions	Not mention ed	26Gy/5F x/1w	N/A	72	Local and regional recurrences	Recruit ing

					M0								
NCT0444 3413	Unite d States	≥18	2020	Yes	BCS or mastectomy; clinical or pathologic T1-T4c, N0-3, M0; indicated for RNI	Not mentioned	Optional Study group: SIB of 5-fraction; Control group: 4-fraction boost of x-ray therapy	Not mention ed	proton therapy over 5 fractions	x-ray therapy over 25 fractions	146	Complication rate	Recruit ing
NCT0422 8991	Cana da	≥18	2021	Yes	BCS or mastectomy; pT3N0, pT1-3N1-2; Or NAC with cT3N0, cT1-3N1-2 and ypT0-3N0-2	No	Not mentioned	Not mention ed	26Gy/5F x/1w	40Gy/15F x/3w	588	Lymphedema	Recruit
NCT0447 2845	India	≥18	2021	Yes	BCS or mastectomy; pT3-4pN2-3 M0; NAC with clinical stage III or ypN+	Tissue expanders with distant metal ports are allowed	Study group: SIB of 34Gy/5Fx or sequential 8Gy/2Fx/2d; Control group: SIB of	for patients with T3-4 central and inner quadran	26Gy/5F x/1w	34Gy/10F x/2w	1018	Locoregional control	Recruit ing

NCT0566 5920	Brazil	≥18	2022	Yes	T3-4N0-3M 0; Or LABC with NAC BCS; pT1-3 and pN1-3a; with indication of	N/A	of 48Gy/15Fx, or sequential 12Gy/4Fx Not mentioned	N2-3	in N+ or after NAC 26Gy/5F x/1w	or after NAC 40Gy/15F x/3w	36	Locoregional recurrence	Recruit
NCT0515 0535	Egypt	≥45	2021	Yes	BCS or mastectomy; TxN1-3M0, T0-2N2-3M 0,	Not mentioned	Study group: SIB of 32Gy/5Fx or sequential 12Gy/4Fx; Control group: SIB	For patients with	26Gy/5F x/1w for WBI/Che st wall, and RNI	40Gy/15F x/3w for WBI/Ches t wall, and RNI in N+	100	Acute and chronic grade 2 toxicity or	Unkno wn
							w or sequential 8Gy/2Fx/2d	and patients with N2					
							42Gy/10fx/2	t lesions					

Abbreviations: WBI = Whole Breast Irradiation; RNI = Regional Nodal Irradiation; IMNI = Internal Mammary Nodal Irradiation; N/A = Not Applicable; SIB = Simultaneous Integrated Boost; w = week; d = days; BCS = Breast-Conserving Surgery; LVSI = Lymphovascular Space Invasion; PBI = Partial Breast Irradiation: NAC = Neoadjuvant Chemotherapy; LABC = Locally Advanced Breast Cancer.