

XXXIII CONGRESSO NAZIONALE AIRO

AIRO2023

BOLOGNA,
27-29 OTTOBRE 2023

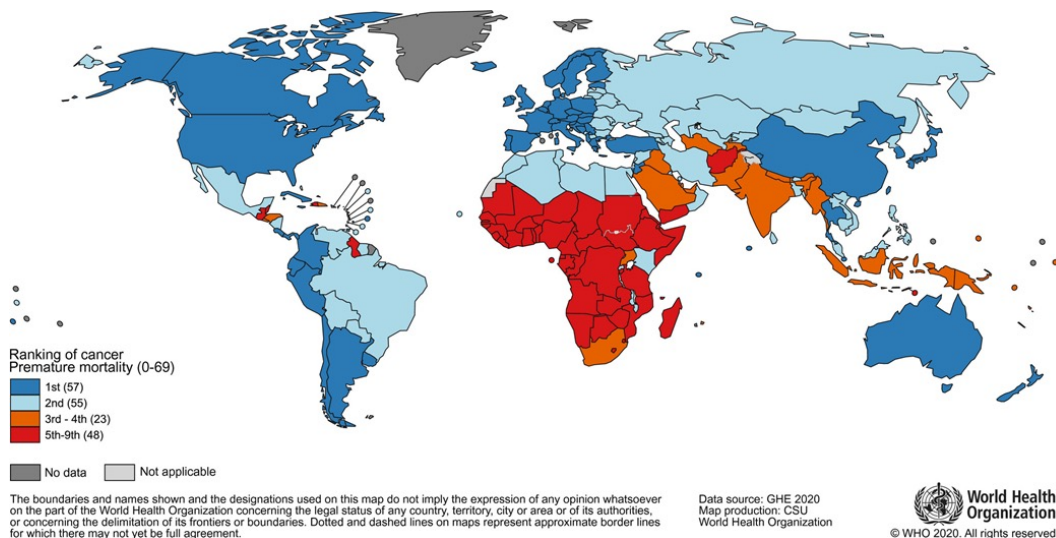
PALAZZO DEI CONGRESSI

Radioterapia Oncologica: l'evoluzione al servizio dei pazienti

INDICAZIONI E TECNICHE DELLA RADIOTERAPIA INTERVENTISTICA NEL TRATTAMENTO DELLE NEOPLASIE MAMMARIE

Antonio Stefanelli
UOC Radioterapia Ferrara

breast cancer world



Background

Over the last 20 years, the therapeutic strategy for breast cancer has increasingly turned towards organ preservation, favoring less mutilating surgical approaches that are constantly integrated with RT and antineoplastic medical therapy.

We have moved from radical mastectomy (aggressive and destructive surgery) to quadrantectomy integrated with post-operative RT (QuaRT, a more conservative and aesthetically friendly treatment) which is currently considered the standard in the early stages of the disease.



BCS + RT vs Mastectomy

	BCS + RT	Mastectomy
LR	5% - 13%	2% - 14%
Survival	59% - 79%	59% - 82%

Milano I, NSABP, EORTC, IGR, NCI, DBCG

Follow up: from 6 to 20 yrs

Hypofractionated EBRT

Hypofractionated breast radiotherapy for 1 week versus 3 weeks (FAST-Forward): 5-year efficacy and late normal tissue effects results from a multicentre, non-inferiority, randomised, phase 3 trial

Adrian Murray Brunt*, Joanne S Haviland[†], Duncan A Whalley, Mark A Sydesham, Abdulla Alhassan, David J Bloomfield, Charlie Chan, Mark Chum, Susan Cleator, Charlotte E Coles, Andrew Goodman, Aditran Hammett, Penelope Hagnwood, Anna M Kirby, Cliona C Kirwan, Carolyn Morris, Zohal Nabi, Elinor Soucy, Navita Somaiah, Liba Stones, Isabel Syrdikus, Judith M Bliss[†], John R Yarnold[†], on behalf of the FAST-Forward Trial Management Group

Published Online

April 28, 2020

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

<https://doi.org/10.1016/j.annonc.2020.04.006>

START Pilot
1986 - 1998

- 42.9Gy in 13Fr
- 39Gy in 13Fr
- 50Gy in 25Fr

START A
1998-2002

- 41.6Gy in 13Fr
- 39Gy in 13Fr
- 50Gy in 25Fr

START B
1999-2001

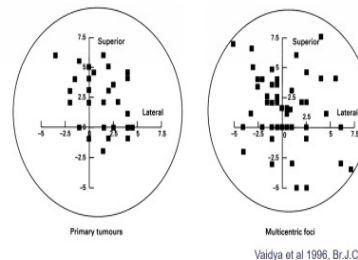
- 40Gy in 15Fr
- 50Gy in 25Fr

Overall there was no significant difference in **loco-regional relapse** at 10 years: approximately 5-8% (for pT1-3 N0-N1)

Moderate of marked breast induration, telangiectasia, breast oedema were significantly less in 39Gy or 40Gy compared to 50Gy

PBI rational

- 1) 70-80% of local recurrences occur in the area of the breast where the primary neoplasm was located
- 2) only 3.3% of them develop a recurrence in an other quadrant than that initially affected by the tumor
- 3) the ipsilateral reappearance of the neoplasm, outside the affected quadrant, is to be considered a second tumor rather than a local recurrence

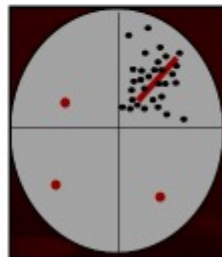


→ 63% of unknown tumor foci outside principal node
→ 80% of unknown foci outside index quadrant

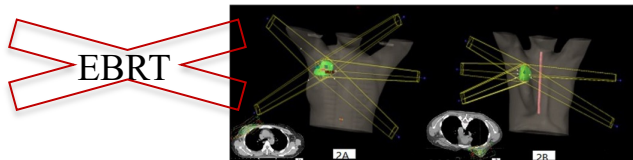
Autors	N° recurrence	% in index quadrant
Clark RM, 1982	680	96%
Schnidt SJ, 1984	231	83%
Boyages J, 1990	783	81%
Kurtz JM, 1990	1593	86%
Fisher B, 1992	1843	100%
Veronesi U, 1993	570	90%

→ About 90% of local failures inside index quadrant

85% of local recurrences occur at or near the original tumor

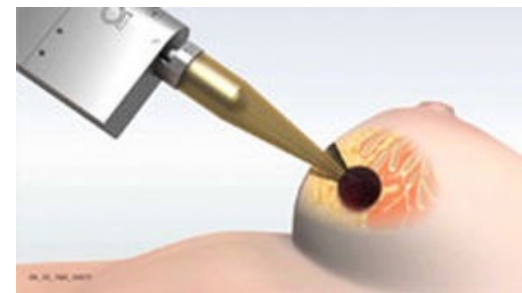


Veronesi U et al. Ann Oncol
2001;12:997-1003.

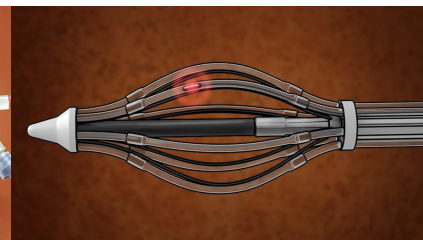
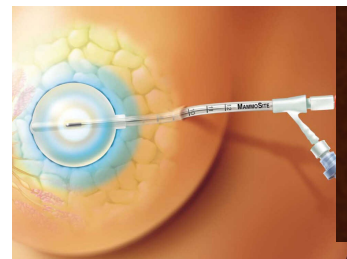


PBI
Partial breast
irradiation
(Boost-
exclusive)

Intra Operative
RadioTherapy (IORT) (E)
KeV (Intrabeam)



Brachitherapy



recurrence

IORT : vantaggi

- ❖ l'esposizione chirurgica permette una accurata definizione del bersaglio
- ❖ l'esposizione chirurgica permette una adeguata protezione dei tessuti sani adiacenti (retrazione tess. mobili, schermatura tess. fissi)
- ❖ buon rapporto efficacia terapeutica / tossicità

IORT TEAM

RADIOTERAPISTA

CHIRURGO

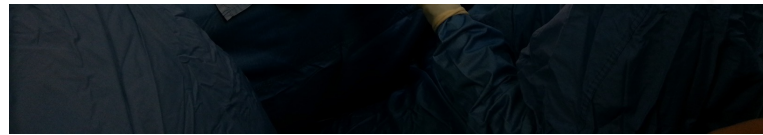
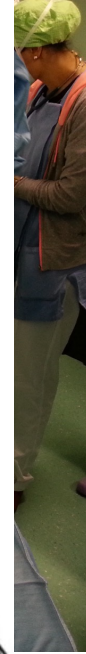
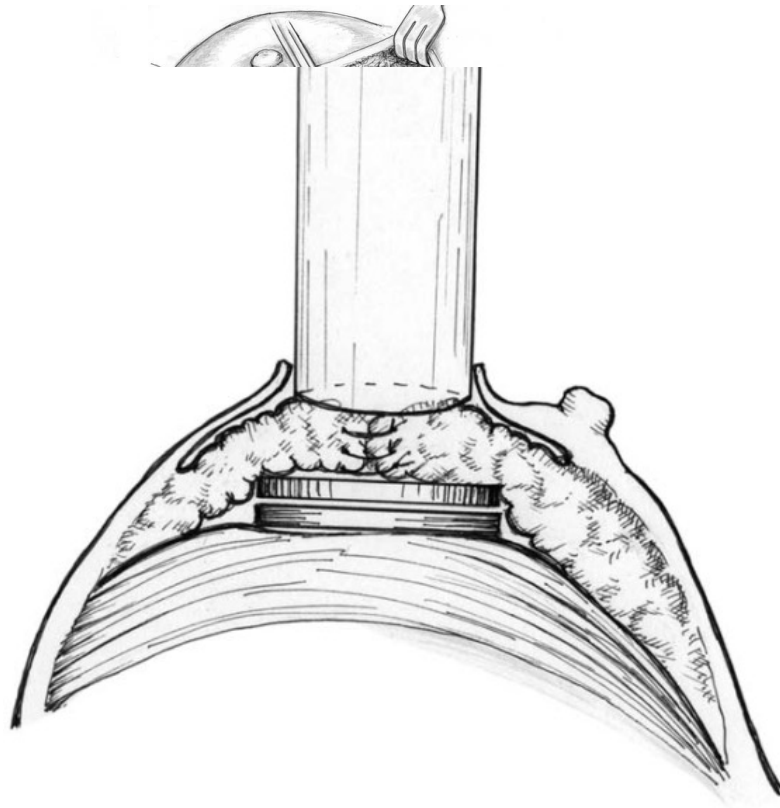
ANESTESISTA

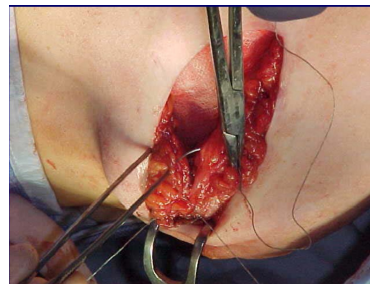
IL FISICO SANITARIO

IL TECNICO DI RADIOLOGIA MEDICA (TSRM)

L'INFERMIERE







Radiation delivery

The image shows a radiation therapy machine (linear accelerator) in a clinical setting. A patient is lying on a treatment table, and the machine's arm is positioned over them. To the right, there is a graph showing the radiation dose distribution. The graph has a vertical axis labeled '100' and a horizontal axis labeled '40', '30', '20', '10', '0', '10', '20', '30', '40'. The graph shows several overlapping curves representing different dose levels. Below the graph, there is a photograph of a breast tumor being treated with radiation, showing the tumor and the surrounding tissue.

AIRO2023

Radioterapia Oncologica:
l'evoluzione al servizio dei pazienti



BOLOGNA, 27-29 OTTOBRE 2023
PALAZZO DEI CONGRESSI



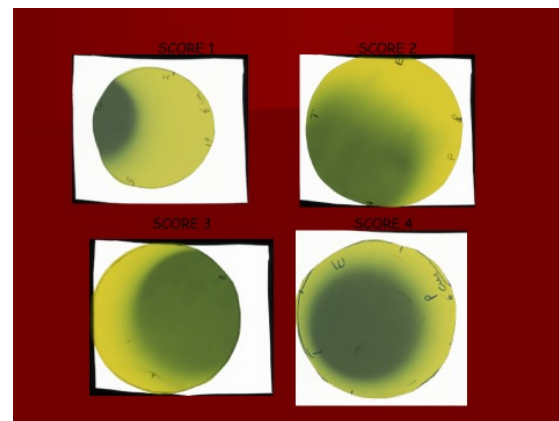


ACCESSORI

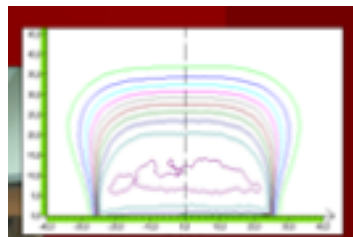
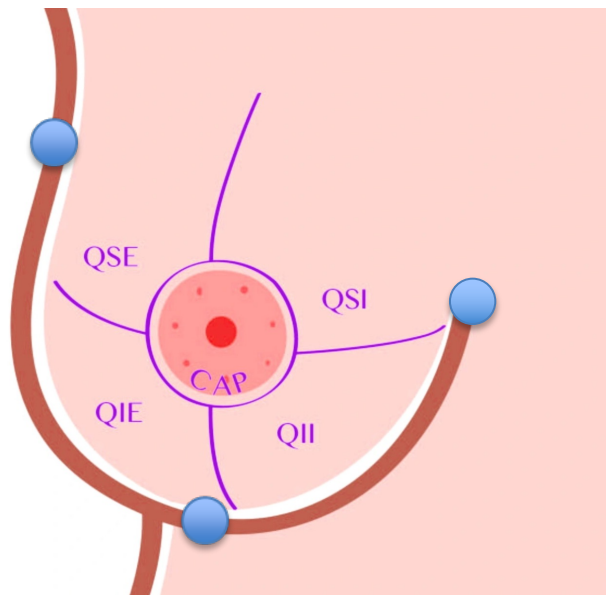
- Applicatori e preapplicatori



- Dispositivi di radioprotezione

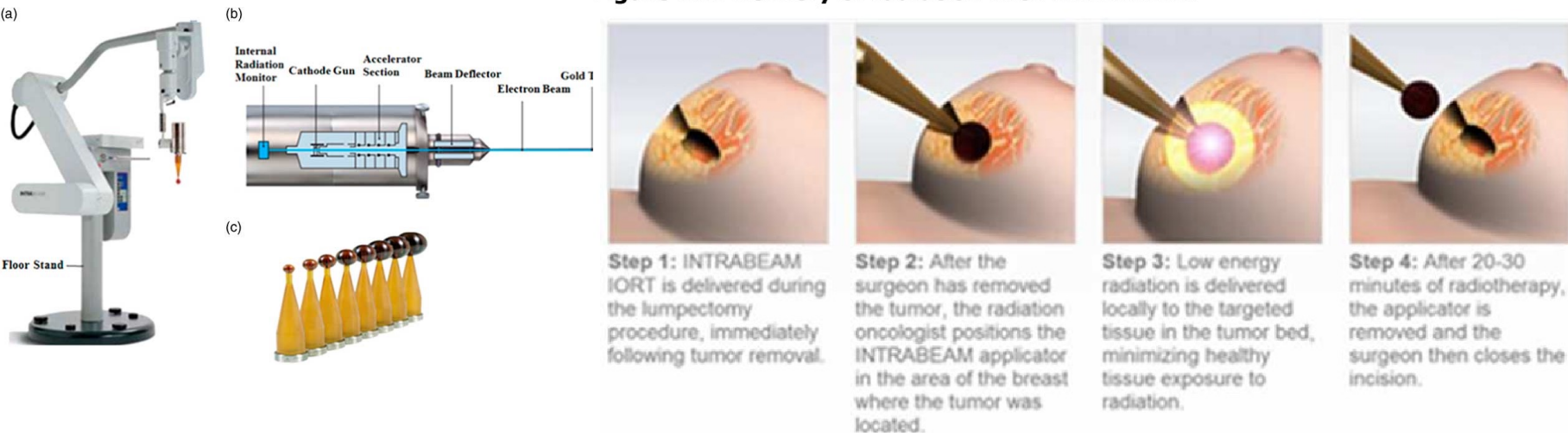


LIMITI ANATOMO-DOSIMETRICI



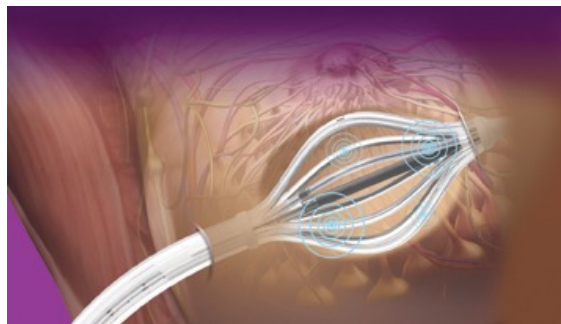
INTRABEAM

Figure 2.2: Delivery of radiation with INTRABEAM



Source: Oncology Systems Limited (2016)

BRACHITERAPIA



AIRO2023

Intraoperative radiotherapy versus external radiotherapy for early breast cancer (ELIOT): a randomised controlled equivalence trial



Umberto Veronesi, Roberto Orecchia, Patrick Maisonneuve, Giuseppe Viale, Nicole Rotmensch, Claudia Sangalli, Alberto Luini, Paolo Veronesi, Viviana Galimberti, Stefano Zurrida, Maria Cristina Leonard, Roberta Lazzari, Federica Cattani, Oreste Gentilini, Mattia Intra, Pietro Caldarella, Bettina Ballardini

Summary

Background Intraoperative radiotherapy with electrons allows the substitution of conventional postoperative whole breast irradiation with one session of radiotherapy with the same equivalent dose during surgery. However, its ability to control for recurrence of local disease required confirmation in a randomised controlled trial.

Methods This study was done at the European Institute of Oncology (Milan, Italy). Women aged 48–75 years with early breast cancer, a maximum tumour diameter of up to 2.5 cm, and suitable for breast-conserving surgery were randomly assigned in a 1:1 ratio (using a random permuted block design, stratified for clinical tumour size [<1.0 cm vs 1.0 – 1.4 cm vs ≥ 1.5 cm]) to receive either whole-breast external radiotherapy or intraoperative radiotherapy with electrons. Study coordinators, clinicians, and patients were aware of the assignment. Patients in the intraoperative radiotherapy group received one dose of 21 Gy to the tumour bed during surgery. Those in the external radiotherapy group received 50 Gy in 25 fractions of 2 Gy, followed by a boost of 10 Gy in five fractions. This was an equivalence trial; the prespecified equivalence margin was local recurrence of 7.5% in the intraoperative radiotherapy group. The primary endpoint was occurrence of ipsilateral breast tumour recurrences (IBTR); overall survival was a secondary outcome. The main analysis was by intention to treat. This trial is registered with ClinicalTrials.gov, number NCT01849133.

Findings 1305 patients were randomised (654 to external radiotherapy and 651 to intraoperative radiotherapy) between Nov 20, 2000, and Dec 27, 2007. After a medium follow-up of 5–8 years (IQR 4.1–7.7), 35 patients in the intraoperative radiotherapy group and four patients in the external radiotherapy group had an IBTR ($p<0.0001$). The 5-year event rate for IBTR was 4.4% (95% CI 2.7–6.1) in the intraoperative radiotherapy group and 0.4% (0.0–1.0) in the external radiotherapy group (hazard ratio 9.3 [95% CI 3.3–26.3]). During the same period, 34 women allocated to intraoperative radiotherapy and 31 to external radiotherapy died ($p=0.59$). 5-year overall survival was 96.8% (95% CI 95.3–98.3) in the intraoperative radiotherapy group and 96.9% (95.5–98.3) in the external radiotherapy group. In patients with data available ($n=464$ for intraoperative radiotherapy; $n=412$ for external radiotherapy) we noted significantly fewer skin side-effects in women in the intraoperative radiotherapy group than in those in the external radiotherapy group ($p=0.0002$).

Published Online
November 11, 2013
[http://dx.doi.org/10.1016/S1470-2045\(13\)70497-2](http://dx.doi.org/10.1016/S1470-2045(13)70497-2)

See Online/Comment
Lancet 2013; published online
November 11,
[http://dx.doi.org/10.1016/S0140-6736\(13\)62304-1](http://dx.doi.org/10.1016/S0140-6736(13)62304-1)

Scientific Directorate
(Prof U Veronesi MD), Division
of Radiation Oncology
(Prof R Orecchia MD,
M C Leonard MD, R Lazzari MD,
F Cattani MD), Division of
Epidemiology and Biostatistics
(P Maisonneuve DipEng,
N Rotmensch MSc), Division of
Pathology (Prof G Viale MD),
and Division of Breast Surgery
(C Sangalli MSc, A Luini MD,
Prof P Veronesi MD,
V Galimberti MD,
Prof S Zurrida MD,
O Gentilini MD, M Intra MD,
P Caldarella MD,
B Ballardini MD), European
Institute of Oncology, Milan,
Italy; and University of Milan,
Milan, Italy (Prof R Orecchia,
Prof G Viale, Prof P Veronesi,
Prof S Zurrida)

	External radiotherapy	Intraoperative radiotherapy with electrons
Age*		
48–49 years	43 (7%)	44 (7%)
50–59 years	267 (41%)	286 (44%)
60–69 years	269 (41%)	259 (40%)
≥70 years	75 (11%)	62 (10%)
Histology†		
Ductal	514 (79%)	524 (81%)
Lobular	57 (9%)	53 (8%)
Ductal and lobular	21 (3%)	17 (3%)
Other	55 (9%)	53 (8%)
Pathological size‡		
≤1 cm	194 (30%)	199 (31%)
1–1.5 cm	235 (36%)	243 (38%)
1.5–2 cm	115 (18%)	120 (19%)
>2 cm	103 (16%)	83 (13%)
Number of positive nodes‡		
None	471 (73%)	478 (74%)
1–3	138 (21%)	138 (21%)
≥4	38 (6%)	31 (5%)
Tumour grade§		
G1	160 (25%)	196 (31%)
G2	328 (52%)	305 (48%)
G3	145 (23%)	129 (20%)
Oestrogen receptor¶		
Negative	56 (9%)	63 (10%)
Positive	589 (91%)	582 (90%)
Progesterone receptor 		
Negative	132 (20%)	158 (24%)
Positive	512 (80%)	487 (76%)
Proliferative index (Ki-67)**		
<14%	242 (38%)	263 (41%)
14–20%	138 (21%)	138 (21%)
>20%	265 (41%)	244 (38%)
Molecular subtype¶¶		
Luminal A	237 (37%)	256 (40%)
Luminal B	352 (55%)	327 (51%)
HER2 positive (non-luminal)	24 (4%)	20 (3%)
Triple negative	32 (5%)	43 (7%)
Adjuvant treatment*		
Control	26 (4%)	25 (4%)
Endocrine therapy alone	485 (74%)	489 (75%)
Chemotherapy alone	47 (7%)	53 (8%)
Endocrine and chemotherapy	96 (15%)	84 (13%)

Data are n (%). Some percentages do not total 100% because of rounding. 654 patients were assigned to external radiotherapy, and 651 to intraoperative radiotherapy with electrons. *n=654 for external radiotherapy, n=651 for intraoperative radiotherapy with electrons. †n=647 for both groups. ‡n=647 for external radiotherapy, n=645 for intraoperative radiotherapy with electrons. §n=633 for external radiotherapy, n=630 for intraoperative radiotherapy with electrons. ¶n=645 for external radiotherapy, n=646 for intraoperative radiotherapy with electrons. ||n=644 for external radiotherapy, n=645 for intraoperative radiotherapy with electrons. **n=645 for both groups.

Table 1: Characteristics of patients according to allocated group (intention-to-treat population)

Radioterapia Oncologica:
l'evoluzione al servizio dei pazienti

AIRO2023

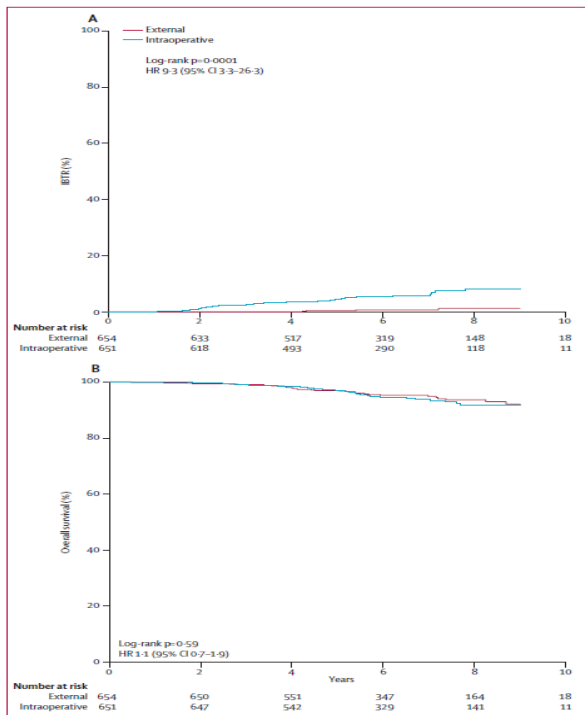


Figure 2: Cumulative incidence of (A) ipsilateral breast tumour recurrence and (B) overall survival

	Patients (n/N)	IBTR 5-year event rate (95% CI)	Log-rank p-value*
Total	35/651	4.4% (2.7-6.1)	--
Age			
48-49 years	0/44	0	--
50-59 years	21/286	5.6% (2.7-8.5)	--
60-69 years	10/259	3.1% (0.8-5.4)	--
≥70 years	4/62	7.2% (0.4-14.1)	0.11
Histology			
Ductal	28/524	4.5% (2.6-6.5)	--
Lobular	3/53	4.6% (0.0-10.8)	--
Ductal and lobular	2/17	6.3% (0.0-18.1)	--
Other	2/53	2.1% (0.0-6.1)	0.69
Pathological size			
≤1 cm	5/199	1.9% (0.0-4.0)	--
1-1.5 cm	13/243	4.2% (1.5-6.9)	--
1.5-2.0 cm	7/120	4.7% (0.7-8.8)	--
>2.0 cm	10/83	10.9% (3.7-18.1)	0.006
Number of positive nodes			
None	21/478	3.5% (1.7-5.3)	--
1-3	10/138	5.3% (1.5-9.2)	--
≥4	4/31	15.0% (1.4-28.7)	0.06
Overall p value	--	--	--
Tumour grade			
G1	5/196	1.1% (0.0-2.7)	--
G2	15/305	3.8% (1.5-6.1)	--
G3	15/129	11.9% (5.7-18.2)	0.0003
Oestrogen receptor			
Absent	8/63	14.9% (5.2-24.5)	--
Present	21/583	3.3% (1.8-4.9)	0.004
Overall p value	--	--	--
Progesterone receptor			
Absent	12/158	7.4% (2.9-11.8)	--
Present	23/487	3.5% (1.7-5.2)	0.17
Proliferative index (Ki-67)			
<14%	8/263	1.8% (0.0-3.5)	--
14-20%	5/138	1.5% (0.0-3.6)	--
>20%	22/244	9.1% (5.3-13.1)	0.002
Molecular subtype			
Luminal A	7/256	1.4% (0.0-3.0)	--
Luminal B	20/327	4.9% (2.4-7.4)	--
HER2-positive (non-luminal)	1/20	5.9% (0.0-17.1)	--
Triple negative	7/43	18.8% (6.1-23.7)	0.001
Characteristics suggesting subsequent whole breast irradiation			
No	14/452	1.5% (0.3-2.7)	--
Yes†	21/199	11.3% (6.4-16.1)	<0.0001

IBTR: ipsilateral breast tumour recurrence. * Overall p value. † Tumour larger than 1 cm, or four or more positive nodes, grade 3, or triple negative.
 Table 3: Factors associated with IBTR among patients randomised to receive intraoperative radiotherapy with electrons

Radioterapia Oncologica: l'evoluzione al servizio dei pazienti

	External radiotherapy	intraoperative radiotherapy with electrons	p value†
Any skin toxicity			
No	427	401	--
Yes, acute	32	5	--
Yes, chronic	5	6	0.0002
Erythema			
No	7	24	--
Grade 1-2	35	5	--
Grade 3	2	0	--
Grade 4	3	0	--
Grade 5	0	0	<0.0001
Dryness			
No	128	147	--
Grade 1-2	20	10	--
Grade 3-5	0	0	0.04
Hyper-pigmentation			
No	138	146	--
Grade 1-2	36	11	--
Grade 3-5	0	0	0.0004
Pruritus (scale 0-10)			
0	174	153	--
1-2	6	5	--
≥3	11	0	0.006
Overall p value	--	--	--
Necrosis (radiological)			
Absent	136	129	--
Present	10	22	0.04

*Information available only for a subset of patients. †Overall p value.

Table 4: Skin side-effects (per-protocol analysis)*

ELIOT should be offered to selected patients at low-risk of IBTR

Taccuino IORT (2014) esclusiva al di fuori di studi clinici: indicazioni

INDICAZIONI

- Età > 50 anni e stato menopausale
- Malattia **unifocale**
- Istotipo invasivo **non** lobulare
- $T \leq 2$ cm
- malattia con profilo biologico favorevole (basso indice di proliferazione, recettori ormonali positivi, HER 2 negativo ovvero gruppo fenotipico **luminal A**)
- assenza di metastasi linfonodali (**NO**)
- margini chirurgici **macroscopicamente** negativi

CONTROINDICAZIONI ASSOLUTE

- Malattia multifocale o multicentrica
- $T > 2$ cm o T4
- N+
- Chemioterapia neoadiuvante
- Mutazione di BRCA
- Gravidanza

CONTROINDICAZIONI RELATIVE

Malattie del connettivo. Alcune malattie del collagene quali lupus, sclerodermia, dermatomiosite se in fase quiescente rappresentano una controindicazione relativa, se in fase attiva assoluta per l'amplificazione delle tossicità segnalate. L'artrite reumatoide non è considerata controindicazione al trattamento

Protocollo di studio IRMA3:

carcinoma della mammella a basso rischio di recidiva locale: irradiazione parziale e accelerata con radioterapia conformazionale tridimensionale (3DCRT) vs. radioterapia standard dopo chirurgia conservativa e radioterapia intraoperatoria (IORT) verso radioterapia standard dopo chirurgia conservativa (studio di fase III).

Popolazione target dello studio

Donne di età \geq 49 anni, ECOG 0-2, sottoposte a chirurgia mammaria conservativa per carcinoma della mammella invasivo, T1-2 (< 2,5 cm di diametro) N0 M0, unifocale, margini di resezione istologicamente negativi (\leq 2 mm) al primo intervento o dopo successivo ampliamento.



Partial breast irradiation *IORT sorgente RX (Intrabeam)*



Caduta estremamente rapida della dose

Trattamento intraoperatorio a ferita aperta dopo verifica margini

Irradiazione della cavità chirurgica a 360°

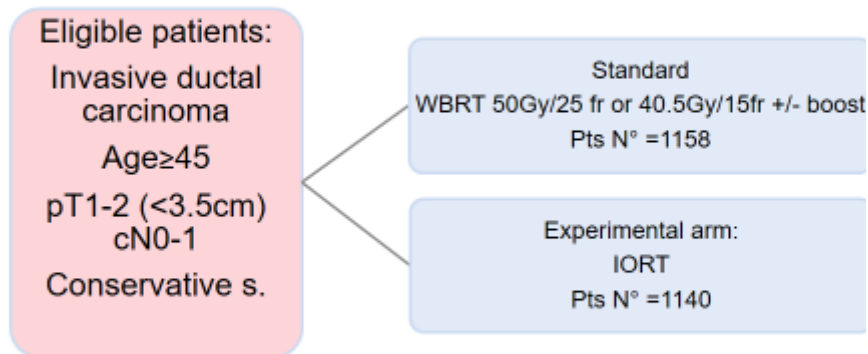
Calcolo della dose a 0.5 cm dalla superficie



Applicator sphere in tumour bed

TARGET –A: trial design

Primary endpoint: local control (non-inferiority)



Median follow up: 8.6 years

BMJ 2020;370:m2836

IORT - PBI - dose

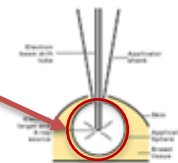
2Gy/fraction equivalent dose of 20 Gy in single fraction:

Dose IORT	α/β	20Gy
Normal tissue (acute tox)	7	60Gy
Tumor	10	50Gy
Normal tissue (late tox)	2	120Gy

RT prescription:

-5Gy at 1cm isodose (=20Gy on applicator surface in water equivalent phantom)

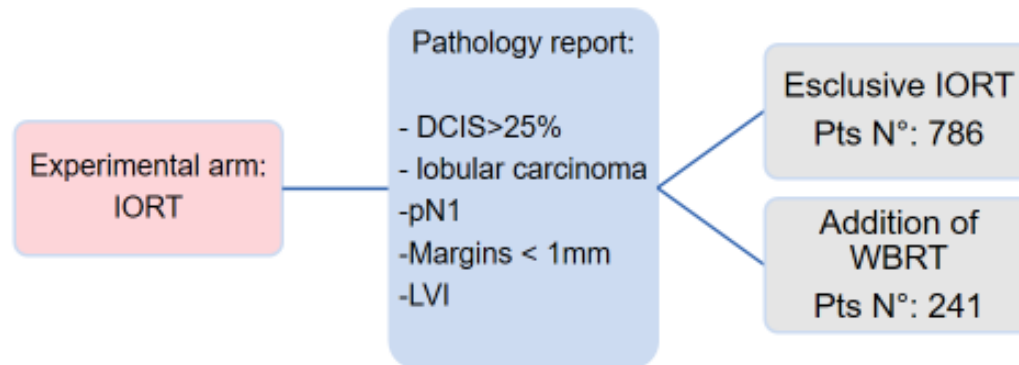
-6Gy at 1 cm isodose (=20Gy on applicator surface in fat equivalent phantom)



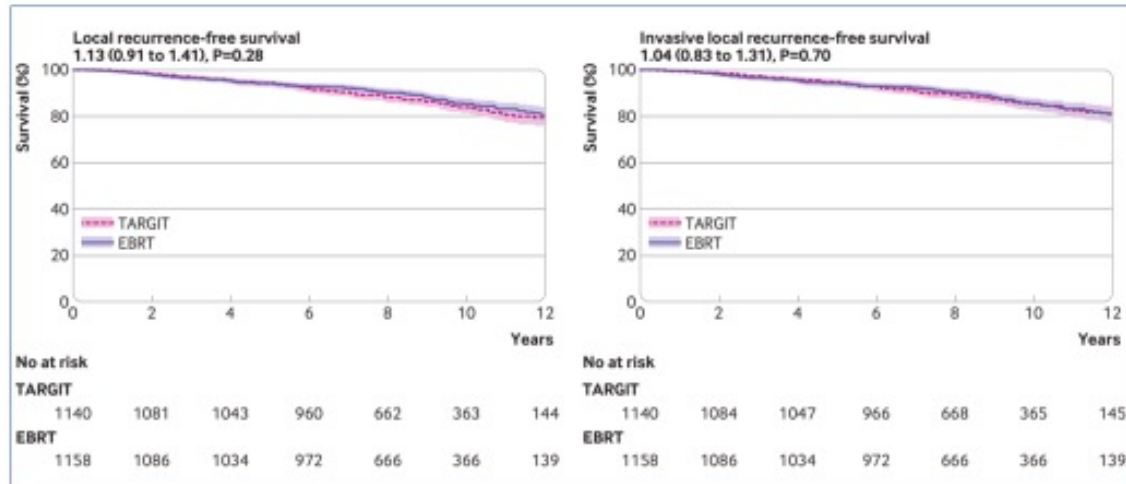
Courtesy Dr. Vinante

Targit-A: trial design

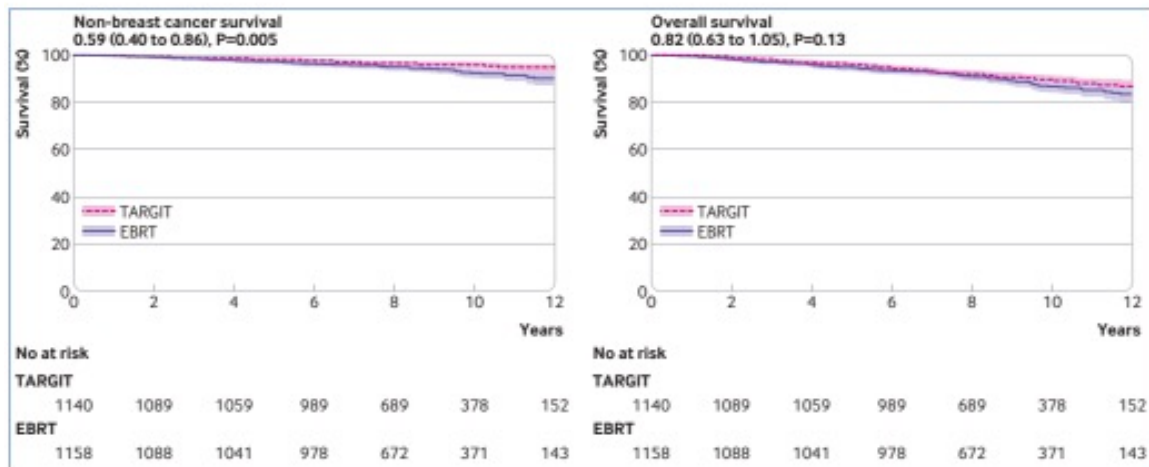
TARGIT – IORT risk adapted treatment:



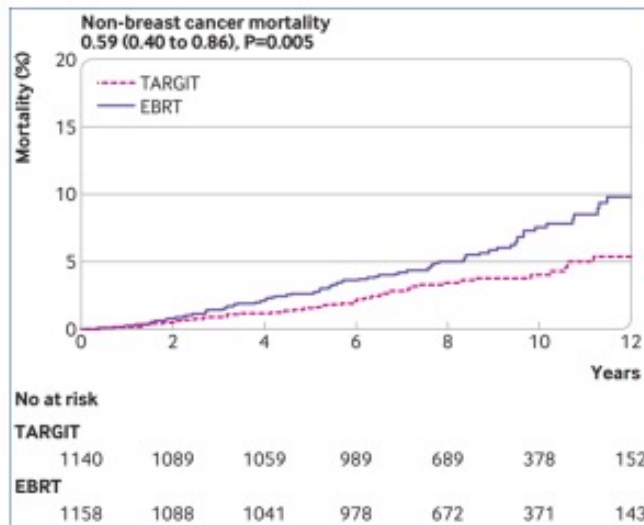
Targit-A: results



Targit-A: results



Targit-A: results



Causes of death	I O R T	E B R T
Breast cancer	65	57
Cardiovascular	8	20
Other cancers	15	21
Other / not given	18	24

BMJ 2020;370:m2836

PBI Brachiterapia

BRACHITERAPIA INTERSTIZIALE



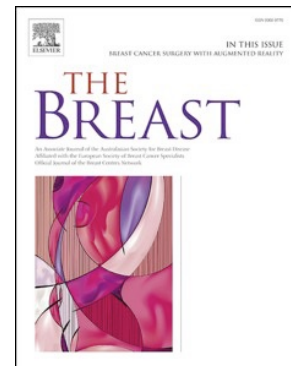
Table 1

Dose and fractionation for brachytherapy or external beam PBI.

RT approach	Total fractionated dose	Dose per fraction	N fractions
Brachytherapy [25,31,35]	30Gy	4.3Gy	7, BID
	32Gy	4Gy	8, BID
	34Gy	3.4Gy	10, BID
External beam radiation therapy [30,31,36–38]	27.5–30Gy	6Gy	5, daily or every other day
	38.5Gy	3.85Gy	10, BID
	40Gy	2.67Gy	15, daily

Abbreviations: Gy, Gray; BID, bis in day.

The Breast 69 (2023) 401–409



Radiotherapy and Oncology 108 (2013) 197–202

Contents lists available at ScienceDirect

Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



ELSEVIER



Phase III randomised trial

Breast-conserving therapy with partial or whole breast irradiation:
Ten-year results of the Budapest randomized trialCsaba Polgár^{a,*}, János Fodor^a, Tibor Major^a, Zoltán Sulyok^b, Miklós Kásler^c^aCenter of Radiotherapy; ^bCenter of Surgery; ^cNational Institute of Oncology, Budapest, Hungary

ARTICLE INFO

Article history:

Received 18 January 2013
Received in revised form 3 May 2013
Accepted 11 May 2013
Available online 3 June 2013

Keywords:

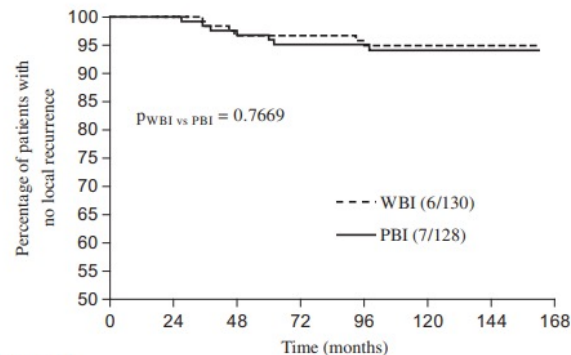
Breast-conserving therapy
Accelerated partial breast irradiation
Brachytherapy
Randomized trial

ABSTRACT

Background and purpose: To report the long-term results of a single-institution randomized study comparing the results of breast-conserving treatment with partial breast irradiation (PBI) or conventional whole breast irradiation (WBI).**Patients and methods:** Between 1998 and 2004, 258 selected women with pT1 pN0-1m1 M0, grade 1–2, non-lobular breast cancer without the presence of extensive intraductal component and resected with negative margins were randomized after 50 Gy WBI (n = 130) or PBI (n = 128). The latter consisted of either 7 × 5.2 Gy high-dose-rate (HDR) multi-catheter brachytherapy (BT; n = 88) or 50 Gy electron beam (EB) irradiation (n = 40). Primary endpoint was local recurrence (LR) as a first event. Secondary endpoints were overall survival (OS), cancer-specific survival (CSS), disease-free survival (DFS), and cosmetic results.**Results:** After a median follow up of 10.2 years, the ten-year actuarial rate of LR was 5.9% and 5.1% in PBI and WBI arms, respectively (p = 0.77). There was no significant difference in the ten-year probability of OS (80% vs 82%), CSS (94% vs 92%), and DFS (85% vs 84%), either. The rate of excellent-good cosmetic result was 81% in the PBI, and 63% in the control group (p < 0.01).**Conclusions:** Partial breast irradiation delivered by interstitial HDR BT or EB for a selected group of early-stage breast cancer patients produces similar ten-year results to those achieved with conventional WBI. Significantly better cosmetic outcome can be achieved with HDR BT implants compared with the outcome after WBI.

© 2013 The Authors. Published by Elsevier Ireland Ltd. Radiotherapy and Oncology 108 (2013) 197–202

Open access under CC BY-NC-ND license.



Number at risk

WBI:	130	128	120	115	111	71	33
PBI:	128	127	122	116	102	63	24

Fig. 2. Time to local recurrence by Kaplan-Meier estimates. *Abbreviations:* WBI – whole breast irradiation; PBI – partial breast irradiation.

Patient selection for PBI

Parameter	GEC-ESTRO	ESTRO-ACROP (2022)
Age	≥50	≥50
histology	NST (no lobular)	NST (no lobular)
T	≤ 3cm and unifocal	≤ 3cm and unifocal
N	N0	N0
Grading	Every	G1-G2
Margins	Negative	Negative > 2mm
Lymphovascular invasion	Absent (focal allowed)	Absent
DCIS	Absent or limited (<25%)	Allowed if < 2.5cm and surgical margins >3mm
Preoperative therapy	none	none
Immunophenotype	every	Luminal A and B

ASTRO consensus statement for APBI

	Suitable (Pt meets <u>all</u> criteria)	Cautionary (Pt meets <u>all</u> criteria)	Unsuitable (Pt meets any criteria)
Age	≥ 50	40-49	< 40
Tumor Size, T stage	≤ 2 cm, Tis or T1	2.1 – 3 cm, T0 or T2	> 3 cm, T3-T4
N stage, surgery	pN0 (SNBx or ALND)		pN1-3 or no nodal surgery
Margins	Negative (≤ 2 mm)	Close (< 2 mm)	Positive
LVSI	No	Limited/focal	Extensive
ER status	Positive	Negative	
Centricity	Unicentric	Microscopic multi-centricity	Present
Histology	Invasive ductal or favorable histology	Invasive lobular	
EIC or Pure DCIS	If screen detected, low to intermediate grade, size ≤2.5 cm, resected with margins negative at >3mm	≤ 3 cm and does not meet criteria for suitable	> 3 cm
Associated LCIS	Allowed		
Neoadjuvant Tx	Not allowed		Received

ASSOCIATION OF RESIDENTS IN RADIATION ONCOLOGY



RAPPORTI ISTISAN 23/5

Quality assurance
in intraoperative radiotherapy in ItalyS. Andreoli, A. Calabrese, C. Di Angelo,
M.C. Lorenzi, L. Meneghini, M. Pignatelli, A. Riva



Clinical Breast Cancer

Volume 22, Issue 2, February 2022, Pages e167-e172



Article



Full-Dose Intraoperative Electron Radiotherapy for Early Breast Cancer: Evidence from a Single Center's Experience

Antonio Stefanelli ^{1,*}, Eleonora Farina ¹, Edoardo Mastella ², Sara Fabbri ², Alessandro Turra ², Simona Bonazza ³, Alessandro De Troia ³, Margherita K. Radica ³ and Paolo Carcoforo ³

LR relapse 5 aa occurred in 2.5% of patients

Clinical Study

Long-Term Outcomes Using Electron IOERT APBI for Early Stage Breast Cancer: The Verona University Hospital Experience

Nunzia Luna Valentina Cernusco ¹  , Paola Del Bianco ², Mario Romano ¹,
Alessandro Muraglia ¹, Gabriella Rossi ¹, Maria Grazia Giri ³, Stefania Guariglia ³,
Davide Lombardi ⁴, Francesca Pellini ⁴, Carlo Cavedon ³, Giovanni Paolo Pollini ⁴,
Renzo Mazzarotto ¹

LR 2.03% 5aa

age, grading, hormone profile.

IORT as reirradiation

Standard treatment for BC recurrence in previously irradiated field: mastectomy

In selected low-risk patients a second lumpectomy is a option

Systematic review of 34 studies of second conservative surgery for BC recurrence:

- 5-year local control : 76% for BCS alone vs. 89% for BCS + re-irradiation
- incidence of grade 3-4 toxicity after re-irradiation: 0 - 21%.
- No phase 3 trials → limited evidence

Walstra CJEF et al. Repeat breast-conserving therapy for ipsilateral breast cancer recurrence: A systematic review. Eur J Surg Oncol. 2019 Aug;45(8):1317-1327. doi: 10.1016/j.ejso.2019.02.008. Epub 2019 Feb 10. PMID: 30795956.

IORT as reirradiation: patient selection

Tumor characteristics:

- Unifocal
 - Unicentric
 - Small size (< 2cm)
- } Mammography + US + MRI

Clinical characteristics:

- Timing first surgery – recurrence: ≥ 24 months
- Adequate breast size

Gentilini O et al. Repeating conservative surgery after ipsilateral breast tumor reappearance: criteria for selecting the best candidates. Ann Surg Oncol. 2012;19(12):3771-8.

IORT as reirradiation

Re-irradiation techniques:

- Brachytherapy
- EBRT (photons, electrons, protons)
- IORT (Intrabeam, electrons)

Authors	N°pts	FUP (m.)	Technique	Dmean	LC (%)	5y-OS	Tox≥G3
Kraus-tiefenbacher	17	26	50kV-X	20	100	-	-
Chin	12	14	50kV-X	20	100	-	0
Thangarajah	41	58	50kV-X	20	89.7	82	0
Blandino	30	47	electrons	18	92.3	94.2	21%

IORT reirradiation series

Montagne L. et al. Second conservative treatment for second ipsilateral breast tumor event: A systematic review of the different re-irradiation techniques. *Breast. 2020;49:274-280.*

AIRO Breast Cancer Group Best Clinical Practice 2022 Update

Re-irradiazione parziale della mammella con brachiterapia

GEC-ESTRO¹² su 217 pazienti ha riportato una rate di terza recidiva, a 5 e 10 anni rispettivamente, del **5.6** e **7.2%**. Nel complesso la tolleranza al trattamento è risultata accettabile.

Lo studio GEC-ESTRO riporta un risultato cosmetico buono/eccellente nell'85% delle pazienti, con una tossicità cutanea e sottocutanea di grado 3 nel 10% dei casi

ASTRO consensus statement for APBI

	Suitable (Pt meets <u>all</u> criteria)	Cautionary (Pt meets <u>all</u> criteria)	Unsuitable (Pt meets <u>any</u> criteria)
Age	≥ 50	40-49	< 40
Tumor Size, T stage	≤ 2 cm, Tis or T1	2.1 – 3 cm, T0 or T2	> 3 cm, T3-T4
N stage, surgery	pN0 (SNBx or ALND)		pN1-3 or no nodal surgery
Margins	Negative (≤ 2 mm)	Close (< 2 mm)	Positive
LVSI	No	Limited/focal	Extensive
ER status	Positive	Negative	
Centricity	Unicentric	Microscopic multi-centricity	Present
Histology	Invasive ductal or favorable histology	Invasive lobular	
EIC or Pure DCIS	If screen detected, low to intermediate grade, size ≤2.5 cm, resected with margins negative at >3mm	≤ 3 cm and does not meet criteria for suitable	> 3 cm
Associated LCIS	Allowed		
Neoadjuvant Tx	Not allowed		Received

ASSOCIATION OF RESIDENTS IN RADIATION ONCOLOGY



CONCLUSION

All the techniques are the same?

Yes in case of adequate selection of patients

Which features allow an adequate selection of patients?

- Histology non-lobular
- Margins preferable > 2 mm
- Biology HR+/HER2-
- Tumor grade preferable G1-2
- Tumor size preferable pT1
- Nodal status preferable pN0 (ITC?)

ONE SHOOT



Article

Hypofractionated Whole Breast Irradiation and Boost-IOERT in Early Stage Breast Cancer (HIOB): First Clinical Results of a Prospective Multicenter Trial (NCT01343459)

Gerd Fastner^{1,*}, Roland Reitsamer², Christoph Gaisberger¹, Wolfgang Hitzl^{3,4,5}, Bartosz Urbański⁶, Dawid Murawa^{7,8}, Christiane Matuschek⁹, Wilfried Budach⁹, Antonella Ciabattini¹⁰, Juliann Reiland^{11,†}, Marie Molnar¹², Cristiana Vidali^{13,†}, Claudia Schumacher¹⁴, Felix Sedlmayer¹ and on behalf of the HIOB Trialist Group[†]

AIRO2023

Radioterapia Oncologica:
l'evoluzione al servizio dei pazienti