

ASSOCIAZIONE ITALIANA RADIOTERAPIA ONCOLOGICA
Piemonte Valle d'Aosta Liguria



Sanremo
13 dicembre 2014

V CONVEGNO
GRUPPO INTERREGIONALE AIRO

Presidente del Convegno: Marco Orsatti

APBI: A CHE PUNTO STIAMO?

MARCO GATTI



QUAL' E' IL RAZIONALE DELLA IRRADIAZIONE PARZIALE DELLA MAMMELLA?

RAZIONALE

- Non è più attuale proporre la stessa radioterapia a tutte le pazienti, soprattutto in un ottica di una moderna personalizzazione del trattamento e del concetto di *“désescalade thérapeutique”*

- L'irradiazione della mammella in toto non è sempre semplice da effettuare sia per motivi di età delle pazienti sia per motivi logistici e socio-economici, al punto che in particolari situazioni il ricorso alla mastectomia è ancora significativo.

RAZIONALE

- La filosofia attuale del trattamento locale del carcinoma della mammella prevede il trattamento dell' intera ghiandola facendo ricorso alla mastectomia totale o associando alla chirurgia conservativa la radioterapia sulla mammella residua.

- Studi istopatologici recenti suggeriscono che nelle pazienti candidate ad una chirurgia conservativa secondo le linee guida attuali sia improbabile l'estensione microscopica di malattia oltre 1 cm dai margini di resezione chirurgica.

Imamura H et al, Breast Cancer Res Treat 2000;62:177–184

Otake T et al., Cancer 1995;76:32–45.

RAZIONALE

- Un altro dato a sostegno della irradiazione parziale della mammella deriva dal fatto che la maggior parte delle recidive locali (85-90%) si verificano nel quadrante mammario dove era localizzata la neoplasia primitiva (*Veronesi et al. Ann Surg Oncol 2001-Malmström et al., Cancer 2003*)
- Il rischio di sviluppare una recidiva omolaterale al di fuori del quadrante interessato è simile a quello di sviluppare un tumore controlaterale (*Veronesi, 2001*) ed è quindi da considerarsi un secondo tumore.
- La RT non diminuisce il tasso di recidiva al di fuori del quadrante interessato

- Alla luce delle nuove acquisizioni si è assistito ad una revisione dello standard terapeutico del trattamento radioterapico del carcinoma della mammella
- Studio di nuove metodiche di irradiazione finalizzate a diminuire la tossicità mediante l'utilizzo di tecniche sofisticate: la radioterapia conformazionale tridimensionale (3D-CRT), la radioterapia a modulazione di intensità (IMRT), la Tomoterapia, la brachiterapia e infine la radioterapia intraoperatoria (IORT)
- Accelerare il trattamento somministrando dosi più alte per frazione consentite dalla diminuzione dei volumi d'irradiazione (APBI)

ACCELERATED PARTIAL BREAST IRRADIATION USING 3D CONFORMAL RADIATION THERAPY (3D-CRT)

KATHY L. BAGLAN, M.D., MICHAEL B. SHARPE, PH.D., DAVID JAFFRAY, PH.D.,
ROBERT C. FRAZIER, M.D., JULIE FAYAD, M.S., LARRY L. KESTIN, M.D.,
VINCENT REMOUCHAMPS, M.D., ALVARO A. MARTINEZ, M.D., F.A.C.R., JOHN WONG, PH.D., AND
FRANK A. VICINI, M.D.

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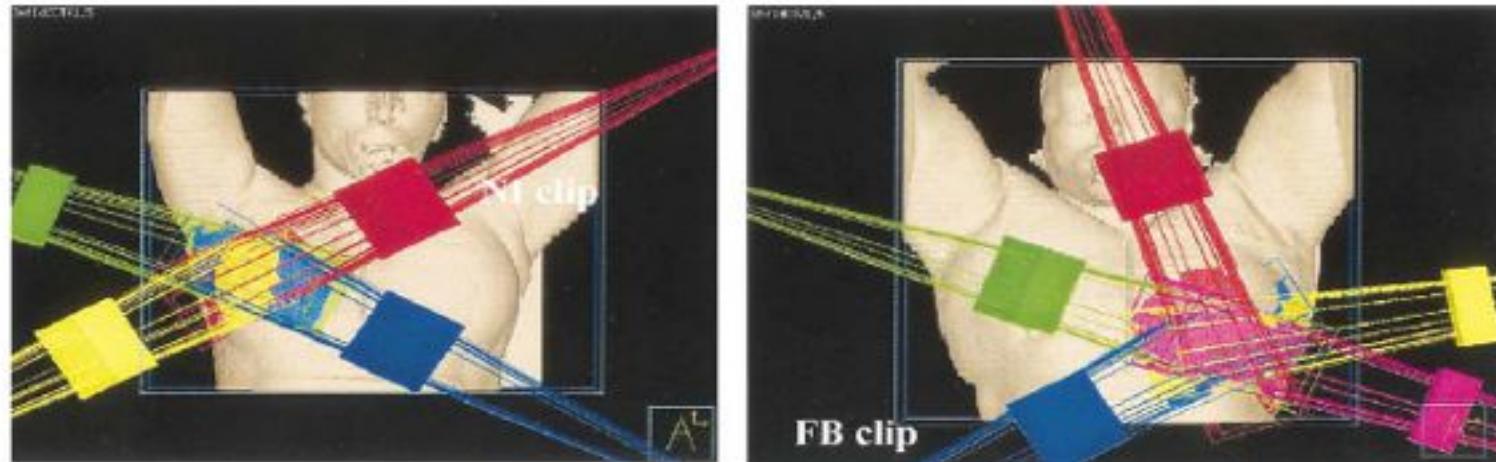
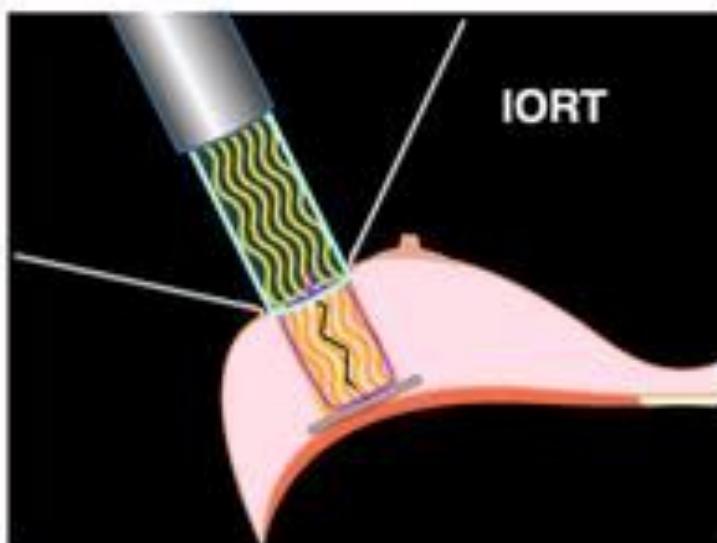
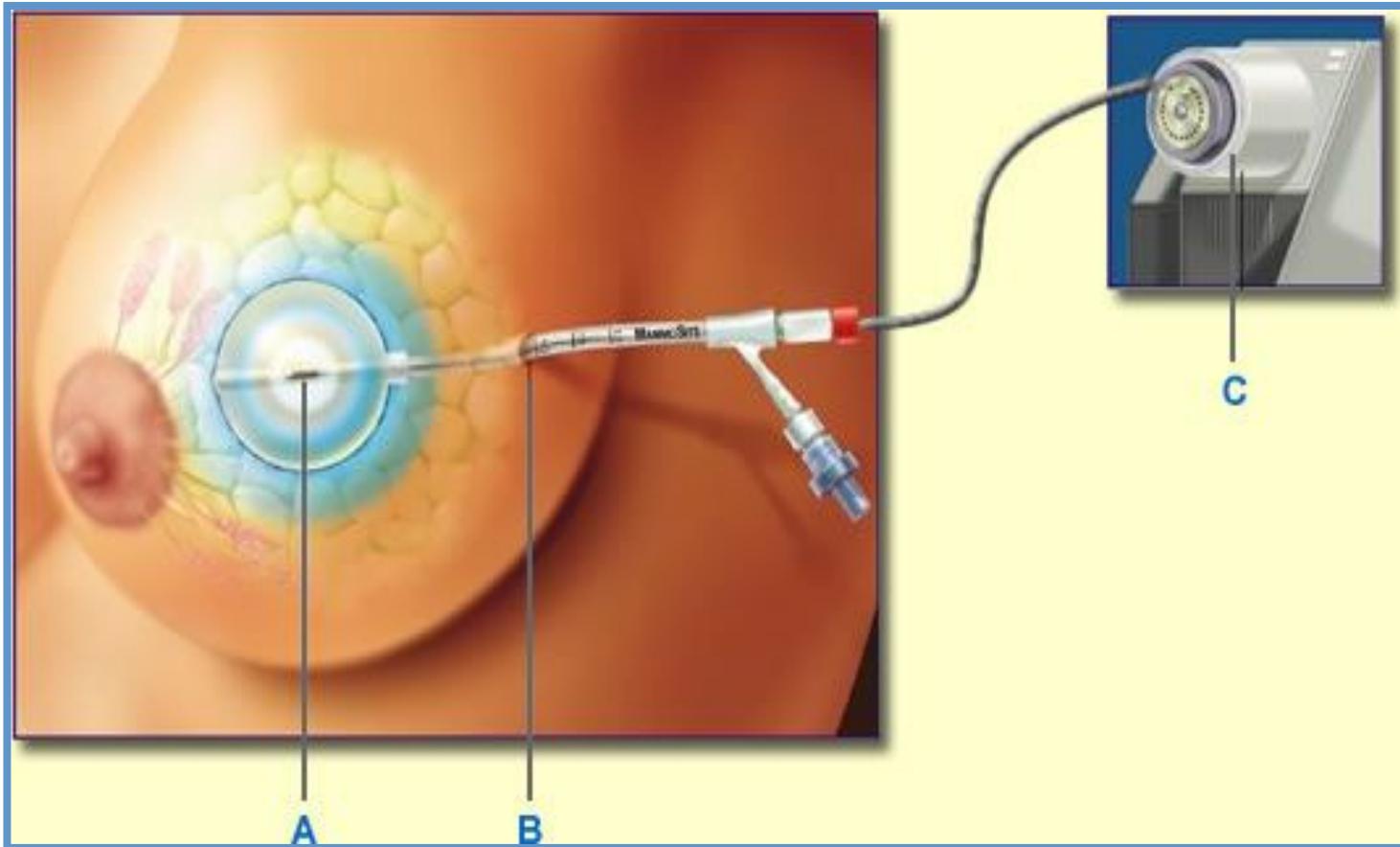


Fig. 1. Typical 4-field arrangement for right-sided lesions and 5-field arrangement for left-sided lesions.

IORT



Mammosite RTS



A small balloon catheter B radioactive seed

C afterloader

IMPLICAZIONI RADIOPATOLOGICHE

- Come si ottiene la dose equivalente a 60 Gy sul letto tumorale mediante l'utilizzo di un frazionamento non convenzionale?
- Il modello lineare quadratico è il più utilizzato:

$$D_1/D_2 = d_2/d_1 + \alpha/\beta / \alpha/\beta$$

D₁= dose totale conv.

D₂= nuova dose equivalente

d₁=dose per frazione conv.

d₂=nuova dose per frazione equivalente

α/β =coeff. resp. lineare e quadratico della “dose dependence”

J Natl Cancer Inst, Vol. 96, No. 3, February 4, 2004

IMPLICAZIONI RADIOPATOLOGICHE

Il rapporto α/β è ricavato empiricamente e nel caso specifico della mammella equivale a circa 4.

- 32 Gy/8 fraz.= 42,7 Gy con $\alpha/\beta=4$
- 34 Gy/10 fraz.= 41,9 Gy con $\alpha/\beta=4$
- 38.5 Gy/10 fraz.=50,4Gy $\alpha/\beta=4$

J Natl Cancer Inst, Vol. 96, No. 3, February 4, 2004

IMPLICAZIONI RADIOPATOLOGICHE

- Le dosi ottenute sono chiaramente inferiori alle dosi utilizzate comunemente nello schema tradizionale, tuttavia la maggior parte degli autori ritengono che una paziente selezionata per PBI e quindi “low risk” richieda una dose inferiore per ottenere un controllo locale equivalente

Bartelink et al., N Engl J Med 2001;345:1378–87.

RESEARCH

Open Access

Higher toxicity with 42 Gy in 10 fractions as a total dose for 3D-conformal accelerated partial breast irradiation: results from a dose escalation phase II trial

Celine Bourgier^{1*}, Catalina Acevedo-Henao¹, Ariane Dunant², Christine Rossier¹, Antonin Levy¹, Mohamed El Nemr^{1,3}, Isabelle Dumas⁴, Suzette Delaloge⁵, Marie-Christine Mathieu⁶, Jean-Remi Garbay⁷, Alphonse Taghian⁸ and Hugo Marsiglia^{1,9}

Conclusions: Early toxicities were more severe and higher rates of late toxicities were observed after 42 Gy/10 fractions/5 days when compared to 40 Gy/10 fractions/5 days. This data suggest that 40 Gy/10 fractions/ 5 days could potentially be the maximum tolerance for PBI although longer follow-up is warranted to better assess late toxicities.

Studi prospettici non randomizzati di fase II

ACCELERATED PARTIAL BREAST IRRADIATION: 5-YEAR RESULTS OF THE GERMAN-AUSTRIAN MULTICENTER PHASE II TRIAL USING INTERSTITIAL MULTICATHETER BRACHYTHERAPY ALONE AFTER BREAST-CONSERVING SURGERY

VRATISLAV STRNAD, M.D.,* GUIDO HILDEBRANDT, M.D.,^{†‡} RICHARD PÖTTER, M.D.,[§]
JOSEF HAMMER, M.D.,[¶] MARION HINDEMITH, M.D.,[†] ALEXANDRA RESCH, M.D.,[§] KURT SPIEGL, M.D.,[¶]
MICHAEL LOTTER, PH.D.,* WOLFGANG UTER, M.D.,^{||} MAYADA BANI,** ROLF-DIETER KORTMANN, M.D.,[†]
MATTHIAS W. BECKMANN, M.D.,** RAINER FIETKAU, M.D.,* AND OLIVER J. OTT, M.D.*

Departments of *Radiation Oncology and **Obstetrics and Gynaecology, University Hospital Erlangen, [†]Department for Medical Informatics, Biometry and Epidemiology, University Erlangen-Nuremberg, Erlangen, [‡]Department of Radiation Oncology, University Hospital Leipzig, Leipzig, ^{||}Department of Radiation Oncology, University Hospital Rostock, Rostock, Germany; [§]Department of Radiation Oncology and Radiobiology, University Hospital AKH Vienna, Vienna, and [¶]Department of Radiation Oncology, Barmherzige Schwestern Hospital, Linz, Austria

274 pts (HDR 32Gy):

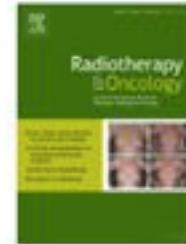
- *Tasso di recidiva locale a 5 aa: 2.9%*
- *Risultato cosmetico: buono/eccellente 92%*
- *Fibrosi grado 3/steatonecrosi: 19,3/4,7 %*



Contents lists available at ScienceDirect

Radiotherapy and Oncology

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Accelerated partial breast irradiation

Accelerated partial-breast irradiation using high-dose-rate interstitial brachytherapy: 12-year update of a prospective clinical study

Csaba Polgár^{a,*}, Tibor Major^a, János Fodor^a, Zoltán Sulyok^b, András Somogyi^a, Katalin Lövey^a, György Németh^a, Miklós Kásler^c

^a Department of Radiotherapy; and ^b Department of General and Thoracic Surgery, National Institute of Oncology, Budapest, Hungary; ^c National Institute of Oncology, Budapest, Hungary

45 pts (HDR 30.3/36.4 Gy):

- *Tasso di recidiva locale a 5/10-12 aa: 4,4/9.3%*
- *Risultato cosmetico: buono/eccezionale 77,8%*
- *Fibrosi grado 3 e steatonecrosi: 2.2%*

FIVE-YEAR ANALYSIS OF TREATMENT EFFICACY AND COSMESIS BY THE
AMERICAN SOCIETY OF BREAST SURGEONS MAMMOSITE BREAST
BRACHYTHERAPY REGISTRY TRIAL IN PATIENTS TREATED WITH ACCELERATED
PARTIAL BREAST IRRADIATION

FRANK VICINI, M.D.,* PETER BEITSCH, M.D.,† CORAL QUIET, M.D.,‡ MARK GITTLEMAN, M.D.,§
VIC ZANNIS, M.D.,|| RICKY FINE, M.D.,¶ PAT WHITWORTH, M.D.,# HENRY KUERER, M.D.,**
BRUCE HAFTY, M.D.,†† MARTIN KEISCH, M.D.,‡‡ AND MAUREEN LYDEN, M.S.|||

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||Department of Surgery, Breast Care Center of the Southwest, Phoenix, AZ; ¶The Breast Center, Marietta, GA; #Nashville Breast Center, Nashville, TN; **Department of Surgery, M.D. Anderson Cancer Center, Houston, TX; ††Robert Wood Johnson University Hospital/Cancer Institute of New Jersey, New Brunswick, NJ; ‡‡Cancer HealthCare Associates, University of Miami Hospital, Miami, FL; and
|||BioStat International, Tampa, FL

1440 pts FU (34 Gy 10 fraz.): 54 mesi

- *Tasso di recidiva locale a 5 aa: 3,04%*
- *Risultato cosmetico: buono/eccezionale 90,6%*
- *Sieromi/steatonecrosi: 13/2.3%*

External Beam Accelerated Partial-Breast Irradiation Using 32 Gy in 8 Twice-Daily Fractions: 5-Year Results of a Prospective Study

Itai M. Pashtan, MD,* Abram Recht, MD,† Marek Ancukiewicz, PhD,‡
Elena Brachtel, MD,§ Rita F. Abi-Raad, MD,‡ Helen A. D'Alessandro, MD,||
Antonin Levy, MD,‡,¶ Jennifer Y. Wo, MD,‡ Ariel E. Hirsch, MD,¶
Lisa A. Kachnic, MD,** Saveli Goldberg, PhD,‡ Michelle Specht, MD,**
Michelle Gadd, MD,** Barbara L. Smith, MD, PhD,** Simon N. Powell, MD, PhD,‡,||
and Alphonse G. Taghian, MD, PhD‡

*Harvard Radiation Oncology Program, Boston, Massachusetts; †Department of Radiation Oncology, Beth Israel Deaconess Medical Center, Boston, Massachusetts; ‡Department of Radiation Oncology, Massachusetts General Hospital, Boston, Massachusetts; §Department of Pathology, Massachusetts General Hospital, Boston, Massachusetts; ¶Department of Radiology, Massachusetts General Hospital, Boston, Massachusetts; ||Department of Radiation Oncology, Boston Medical Center, Boston University School of Medicine, Boston, Massachusetts; **Department of Surgery, Massachusetts General Hospital, Boston, Massachusetts

98 pts (32 Gy 8 fraz.) FU: 71 mesi

- Tasso di recidiva locale a 5 aa: 5%
- Risultati favorevoli nei pz. ER+
- Risultati sfavorevoli nei pz “triple negative”

CONSENSUS STATEMENT

ACCELERATED PARTIAL BREAST IRRADIATION CONSENSUS STATEMENT FROM THE AMERICAN SOCIETY FOR RADIATION ONCOLOGY (ASTRO)

Table 2. Patients "suitable" for APBI if all criteria are present

Factor	Criterion
Patient factors	
Age	≥60 y
BRCA1/2 mutation	Not present
Pathologic factors	
Tumor size	≤2 cm*
T stage	T1
Margins	Negative by at least 2 mm
Grade	Any
LVSI	No†
ER status	Positive
Multicentricity	Unicentric only
Multifocality	Clinically unifocal with total size ≤2.0 cm‡
Histology	Invasive ductal or other favorable subtypes§
Pure DCIS	Not allowed
EIC	Not allowed
Associated LCIS	Allowed
Nodal factors	
N stage	pN0 (i ⁻ , i ⁺)
Nodal surgery	SN Bx or ALND
Treatment factors	
Neoadjuvant therapy	Not allowed

Table 3. "Cautionary" group: Any of these criteria should invoke caution and concern when considering APBI

Factor	Criterion
Patient factors	
Age	50–59 y
Pathologic factors	
Tumor size	2.1–3.0 cm*
T stage	T0 or T2
Margins	Close (<2 mm) Limited/focal
LVSI	Negative†
ER status	Multifocality
Multifocality	Clinically unifocal with total size 2.1–3.0 cm‡
Histology	Invasive lobular
Pure DCIS	≤3 cm
EIC	≤3 cm

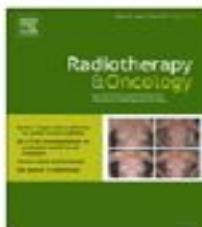
CONSENSUS STATEMENT

ACCELERATED PARTIAL BREAST IRRADIATION CONSENSUS STATEMENT FROM THE AMERICAN SOCIETY FOR RADIATION ONCOLOGY (ASTRO)

BENJAMIN D. SMITH, M.D., *† DOUGLAS W. ARTHUR, M.D., ‡ THOMAS A. BUCHHOLZ, M.D., †
BRUCE G. HAFTY, M.D., § CAROL A. HAHN, M.D., || PATRICIA H. HARDENBERGH, M.D., ¶
THOMAS B. JULIAN, M.D., # LAWRENCE B. MARKS, M.D., ** DORIN A. TODOR, PH.D., ‡
FRANK A. VICINI, M.D., †† TIMOTHY J. WHELAN, M.D., ‡‡ JULIA WHITE, M.D., §§ JENNIFER Y. WO, M.D., |||
AND JAY R. HARRIS, M.D. ¶¶

Table 4. Patients “unsuitable” for APBI outside of a clinical trial if any of these criteria are present

Factor	Criterion
Patient factors	
Age	<50 y
BRCA1/2 mutation	Present
Pathologic factors	
Tumor size*	>3 cm
T stage	T3-4
Margins	Positive
LVSI	Extensive
Multicentricity	Present
Multifocality	If microscopically multifocal >3 cm in total size or if clinically multifocal
Pure DCIS	If >3 cm in size
EIC	If >3 cm in size
Nodal factors	
N stage	pN1, pN2, pN3
Nodal surgery	None performed
Treatment factors	
Neoadjuvant therapy	If used



GEC-ESTRO Recommendations

Patient selection for accelerated partial-breast irradiation (APBI) after breast-conserving surgery: Recommendations of the Groupe Européen de Curiethérapie-European Society for Therapeutic Radiology and Oncology (GEC-ESTRO) breast cancer working group based on clinical evidence (2009)

GEC-ESTRO recommendations on patient selection for accelerated partial-breast irradiation.

Characteristic	A/low-risk group - good candidates for APBI	B/intermediate-risk group - possible candidates for APBI	C/high-risk group - contraindication for APBI
Patient age	>50 years	>40–50 years	≤40 years
Histology	IDC, mucinous, tubular, medullary, and colloid cc.	IDC, ILC, mucinous, tubular, medullary, and colloid cc	-
ILC	Not allowed	Allowed	-
Associated LCIS	Allowed	Allowed	-
DCIS	Not allowed	Allowed	-
HG	Any	Any	-
Tumour size	pT1–2 (≤ 30 mm)	pT1–2 (≤ 30 mm)	pT2 (>30 mm), pT3, pT4
Surgical margins	Negative (≥ 2 mm)	Negative, but close (<2 mm)	Positive
Multicentricity	Unicentric	Unicentric	Multicentric
Multifocality	Unifocal	Multifocal (limited within 2 cm of the index lesion)	Multifocal (>2 cm from the index lesion)
EIC	Not allowed	Not allowed	Present
LVI	Not allowed	Not allowed	Present
ER, PR status	Any	Any	-
Nodal status	pN0 (by SLNB or ALND ^a)	pN1mi, pN1a (by ALND ^a)	pNx; $\geq pN2a$ (4 or more positive nodes)
Neoadjuvant chemotherapy	Not allowed	Not allowed	If used

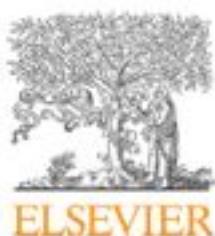
Original Study

Five-Year Outcomes and Toxicities Using 3-Dimensional Conformal External Beam Radiation Therapy to Deliver Accelerated Partial Breast Irradiation

Chirag Shah,¹ J. Ben Wilkinson,² Thomas Lanni,² Maha Jawad,² Jessica Wobb,²
Ashley Fowler,² Michelle Wallace,² Peter Chen,² Inga S. Grills,² Frank Vicini³

192 pts (34/38.5 Gy 10 fraz.): 37,5% suitable, 46,4% cautionary, 16,1% insuitable according to ASTRO

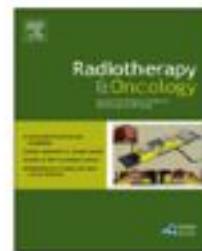
- *Tasso di recidiva locale a 5 aa: 0%*
- *Risultato cosmetico: buono/eccezionale 81%*
- *Fibrosi grado 3: 7.6%*
- *Teleangectasie: 7.5%*



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Accelerated partial breast irradiation

Defining an optimal role for breast magnetic resonance imaging when evaluating patients otherwise eligible for accelerated partial breast irradiation



Kathleen C. Horst^{a,*}, Katherine E. Fero^a, Debra M. Ikeda^b, Bruce L. Daniel^b, Frederick M. Dirbas^c

^aDepartment of Radiation Oncology; ^bDepartment of Radiology; ^cDepartment of Surgery, Stanford University School of Medicine, Stanford Cancer Institute, United States

- *La RM è stata in grado di identificare una malattia multifocale/multicentrica risp. nel 10.5 e nell' 1,6% dei 181 pazienti dello studio inizialmente candidati alla PBI con la stadiazione convenzionale*
- *L'età pre-menopausale e i tumori >2 cm sono i fattori di rischio maggiori*

Studi prospettici randomizzati di fase III

Clin Oncol (R Coll Radiol). 1993;5(5):278-83.

**The Christie Hospital breast conservation trial:
an update at 8 years from inception.**

Ribeiro GG1, Magee B, Swindell R, Harris M,
Banerjee SS.

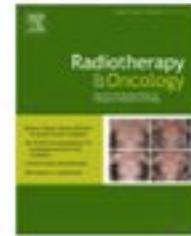
- 708 pts FU 65 mesi
- 40/42.5 Gy/8 fraz./electroni vs whole breast
- Recidive locali 11% vs 15% nel CDI
- Recidive locali: 8% vs 34 % nel CLI
- Sopravvivenza sovrapponibile: 71.2 vs 72,7 %



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Accelerated partial breast irradiation

Accelerated partial-breast irradiation using high-dose-rate interstitial brachytherapy: 12-year update of a prospective clinical study

Csaba Polgár ^{a,*}, Tibor Major ^a, János Fodor ^a, Zoltán Sulyok ^b, András Somogyi ^a, Katalin Lövey ^a, György Németh ^a, Miklós Kásler ^c

^a Department of Radiotherapy; and ^b Department of General and Thoracic Surgery, National Institute of Oncology, Budapest, Hungary; ^c National Institute of Oncology, Budapest, Hungary

- **258 pts selez.+++:** *50 Gy/25 fraz whole breast vs APBI HDR 36,4 Gy/7 fraz o elettroni 6-15 Mev 50 Gy/25 fraz.*
- *Tasso di recidiva locale 5.1 vs 5.9 %*
- *DFS e OS: sovrapponibili*
- *Risultato cosmetico 81% buono/eccezionale APBI vs 63% whole breast*
- *Scarso n. paz, tecniche APBI non confrontabili*

Études randomisées de phase III rapportant les résultats d'efficacité en termes de contrôle local et de survie.

Études	n	Critères d'inclusion	Suivi médian (années)	Protocoles d'irradiation	Résultats		
					Taux de récidives locales (%)	Taux de survie globale (%)	
Budapest [47]	258	> 40 ans T1 unifocal non lobulaire grade 1-2 sans composante intracanalaire extensive Marges chirurgicales saines pN0-1(mi)	5,5	ITS	50 Gy/25 fractions	3	91,8
TARGIT-A [49]	2451	≥ 45 ans T1-2 unifocal non lobulaire sans composante intracanalaire extensive Marges chirurgicales saines pN0-1	2,5	IPAS	HDD 36,4 Gy/7 fractions γ/e- 50 Gy/25 fractions 50 Gy/25 fractions ± boost	4,7	94,6
ELIOT [50]	1305	48-75 ans T1-2 < 2,5 cm unicentrique pNO	5,8	ITS	IORT 50 kV 20 Gy/1 fraction 50 Gy/25 fractions ± 10 Gy	3,3	96,1
				IPAS	IORT e- 21 Gy/1 fraction	0,4	96,9
						4,4	96,8

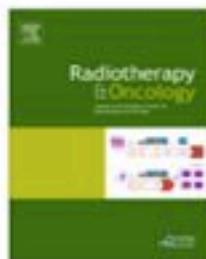
ITS : irradiation de la totalité du sein ; IPAS : irradiation partielle et accélérée du sein ; HDD : curiethérapie de haut débit de dose ; γ : photons gamma de télécobalt ; e- : électrons ; IORT : irradiation intra-opératoire.

Études randomisées de phase III rapportant les résultats de toxicité et les résultats cosmétiques.

Études	n	Critères d'inclusion	Suivi médian (années)	Protocoles d'irradiation	Toxicité			
					Grade 1 (%)	Grade 2 (%)	Grade 3 (%)	
Gec-ESTRO [54]	1193	≥ 40 ans T1-2 ≤ 3 cm pN0-1(mi) marges chirurgicales ≥ 2 mm	1	ITS	50-50,4 \pm 10 Gy	Épidermite : 49,5	-	Épidermite : 7,1
						Hématome : 1,8	-	Hématome : 0,8
				IPAS	HDD 32-30,3 Gy PDR 50 Gy	Atrophie cutanée : 72,3 Hyperpigmentation : 72,7	Atrophie cutanée : 3,7 Hyperpigmentation : 0,65	-
						Épidermite : 18,2 Hématome : 18,8	-	Épidermite : 0,2 Hématome : 0,6
NSABP B39 [56]	4300	≥ 18 ans CCI/CCIS T1-2 ≤ 3 cm $\leq 3N^+$ Marges chirurgicales > 0	3,4	ITS	50-50,4 \pm 10-16 Gy	Atrophie cutanée : 77,8 Hyperpigmentation : 78,5 -	Atrophie cutanée : 1,1 Hyperpigmentation : 0,65 Fibrose ≤ 12	Fibrose ≤ 3
RAPID [57] ^a	2135	≥ 40 ans CCI/CCIS T1-2 ≤ 3 cm pN0 Marges chirurgicales > 0 non BRCA1/2	3	ITS	IPAS RC3D 38,5 Gy 50 \pm 10 42,5 Gy	3	17 ^a	30
				IPAS	RC3D 38,5 Gy	12		

n : nombre de patients ; CCI : carcinome canalaire invasif ; CCIS : carcinome canalaire in situ ; ITS : irradiation de la totalité du sein ; IPAS : irradiation partielle et accélérée du sein ; HDD : curiethérapie de haut débit de dose ; PDR : pulsed dose rate ; RC3D : radiothérapie externe conformatrice tridimensionnelle.

^a Évaluation cosmétique faite pour l'ITS : par l'infirmier 17 %/la patiente 18 %/l'oncologue 17 % (moyenne = 17 %). Évaluation cosmétique faite pour l'IPAS : par l'infirmier 29 %/la patiente 26 %/l'oncologue 35 % (moyenne 30 %).



Review

Accelerated partial irradiation for breast cancer: Systematic review and meta-analysis of 8653 women in eight randomized trials

Gustavo Nader Marta^{a,b,*}, Cristiane Rufino Macedo^d, Heloisa de Andrade Carvalho^{a,c}, Samir Abdallah Hanna^a, João Luis Fernandes da Silva^a, Rachel Riera^d

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^d Brazilian Cochrane Center and Discipline of Emergency Medicine and Evidence-Based Medicine, Universidade Federal de São Paulo-Escola Paulista de Medicina (UNIFESP-EPM), Brazil

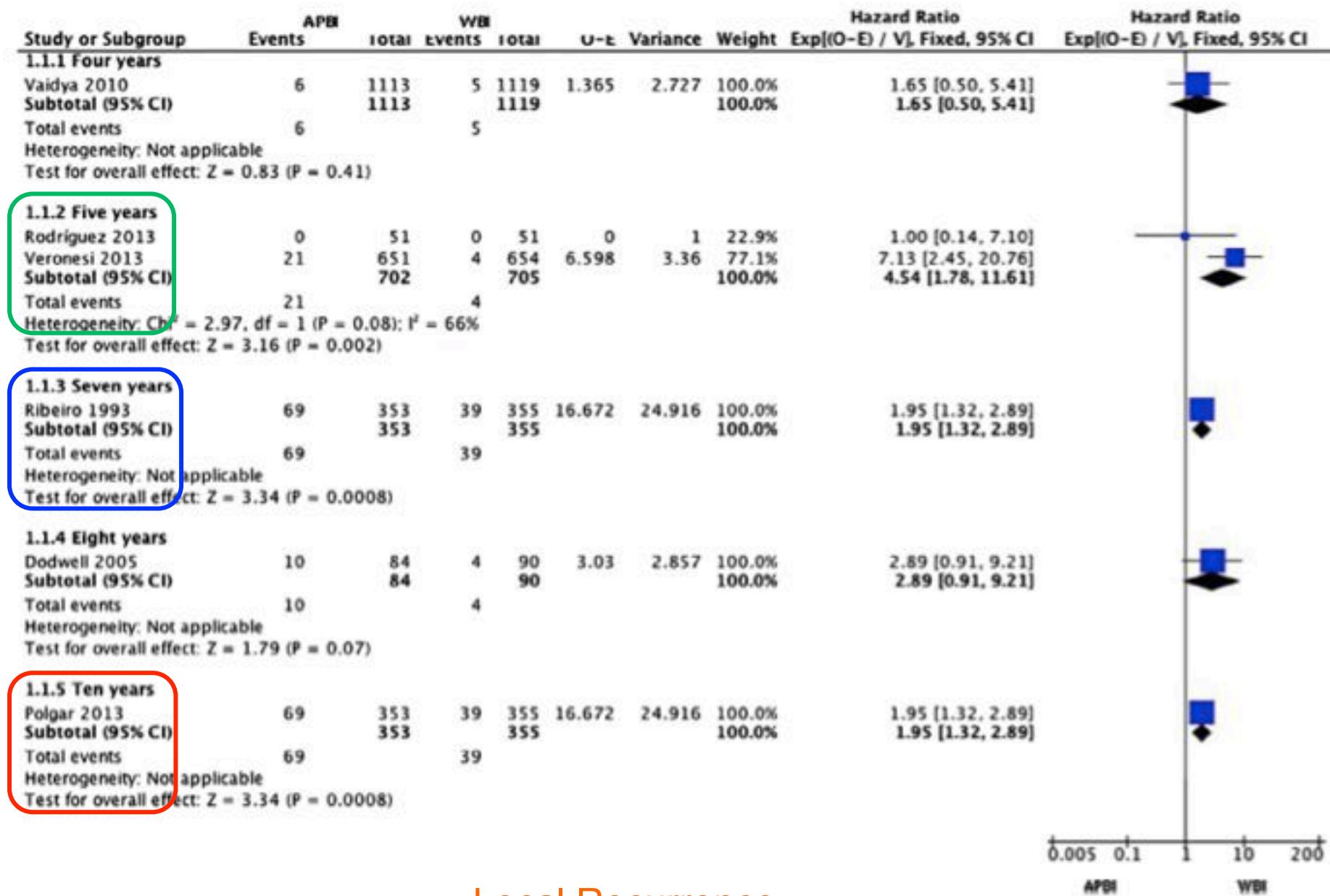
Characteristics of included studies.

Study	Patients (n)	Years of inclusion	Inclusion criteria	Tumor stage (n)			Nodal stage (n)			Grade (n)		ER+ (n)	HER-2+ (n)	WBI arm	APBI arm	Surgery	Che (n)		HT (n)	
				Tis	T1-T2	T3-T4	N0	N1-N2	N3	I-II	III						WBI	APBI	WBI	APBI
Dodwell	174	1985-1990	pT1-2 pN0-1	0	174	0	-	WBI 31%	APBI 41%	133	41	-	-	40 Gy/15fx + boost (15 Gy/5fx)	55 Gy/20fx	Local excision and a level 2 ALND	90	84	90	84
Livi	520	2005-2008	>40 years; T ≤ 2.5 cm; unifocal tumor; no EIC	39	220	0	206	48	11	201	29	202 [†]	-	50 Gy/25fx	30 Gy/5fx (IMRT)	Wide excision or quadrantectomy, clear margins (>5 mm)	8	7	73	68
Olivetto	2135	2006-2011	>40 years; DCIS; IDC < 3 cm; pN0	366	1752	0	2135	0	0	1425	302	1471	-	50 Gy/25fx (large breast size) or 42.5 Gy/16fx +/- boost	38.5 Gy/10fx twice daily	BCS, negative margins, negative SLNB or ALND	137	130	582	604
Polgar	258	1998-2004	Unifocal tumor; <pT2; cN0, pN0, or pN1mi (single nodal >0.2 mm and <2 mm); histologic grade <2	0	258	0	249	9	0	258	0	229	-	40-50 Gy/21-25fx	HDR MI 36.4 Gy/7fx or electrons 42-50 Gy/21-25fx	Wide excision, negative margins, ALND/SLNB in some cases	2	2	87	89
Ribeiro	708	1982-1987	<70 years; tumor <4 cm; cN0	-	708	0	-	-	-	-	-	-	40 Gy/15fx (breast and regional nodes)	40-42.5 Gy/8fx (electrons)	Lumpectomy, no ALND	0	0	0	0	
Rodriguez	102	-	>60 years; IDC; unifocal tumor; <pT2; cN0 (pN0 axillary status); histologic grade <2	0	102	102	0	0	0	102	0	100	1	48 Gy/24fx +/- boost (10 Gy/5fx)	37.5 Gy/10fx twice daily	BCS	2	1	51	50
Veronesi	1305	2000-2007	48-75 years; tumor <2.5 cm	0	1305	0	949	276	69	989	274	1172	44	50 Gy/25fx + boost (10 Gy/5fx)	21 Gy/1fx (intraoperative electrons)	Quadrantectomy and SLNB or ALND	47	53	485	489
Vaidya	3451	2000-2012	>45 years; IDC; T1-T2; unifocal tumor; no EIC	0	3451	0	1764	304	61	1769	319	1943	264	40-56 Gy +/- boost 16 Gy	20 Gy (surface of the tumor bed)/5-7 Gy at 1 cm depth	Wide excision	141	116	753	727
Total	8653	-	-	405	7970	102	5303	637	141	4877	965	4915	309	-	-	-	427	393	2121	2111

Note: ALND - Axillary lymph node dissection; BCS - Breast-conserving surgery; Che - Adjuvant therapy with chemotherapy; DCIS - Ductal carcinoma in situ; EIC - Extensive intraductal carcinoma; ER+ - Estrogen receptors positive; HDR MI - High-dose-rate multicatheter interstitial; HER-2+ - HER-2 positive; HT - Adjuvant therapy with hormone; IDC - Invasive ductal carcinoma; IMRT - Intensity-modulated radiotherapy; LVS - Lymphovascular space invasion; SLNB - Sentinel lymph node biopsy.

[†] Hormonal receptors positive.

Accelerated partial breast irradiation: Meta-analysis



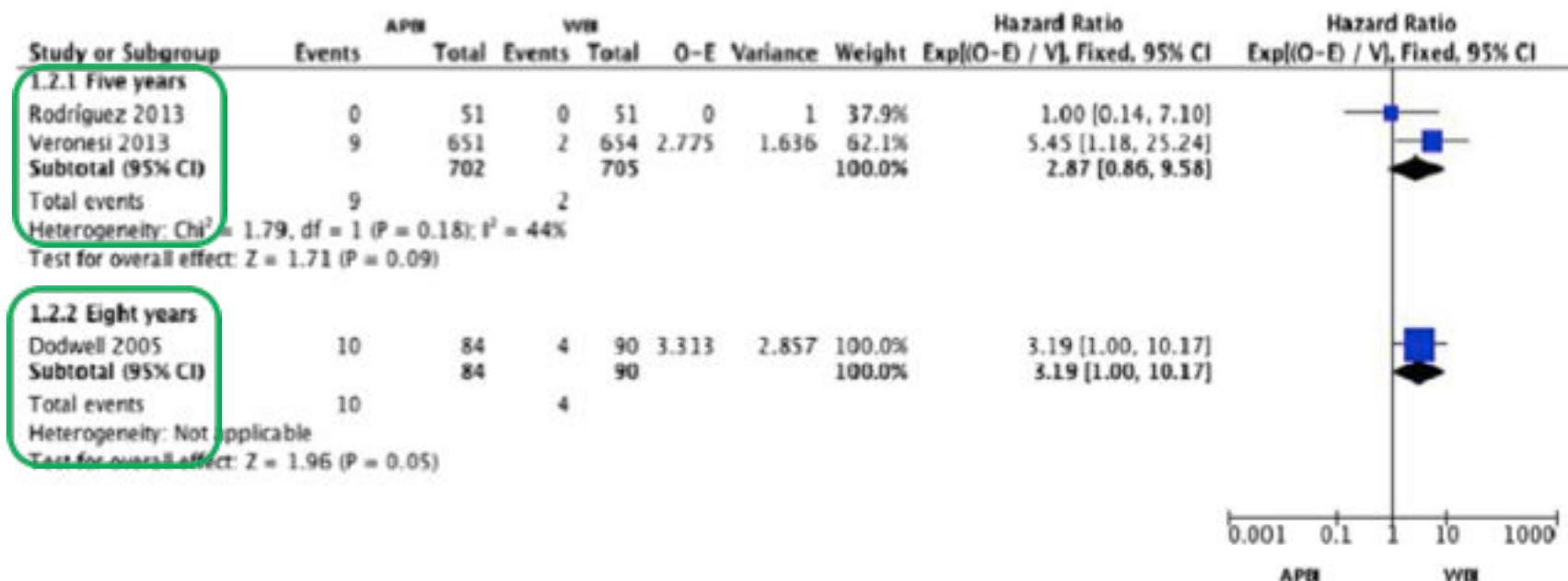


Fig. 2. Nodal recurrence.

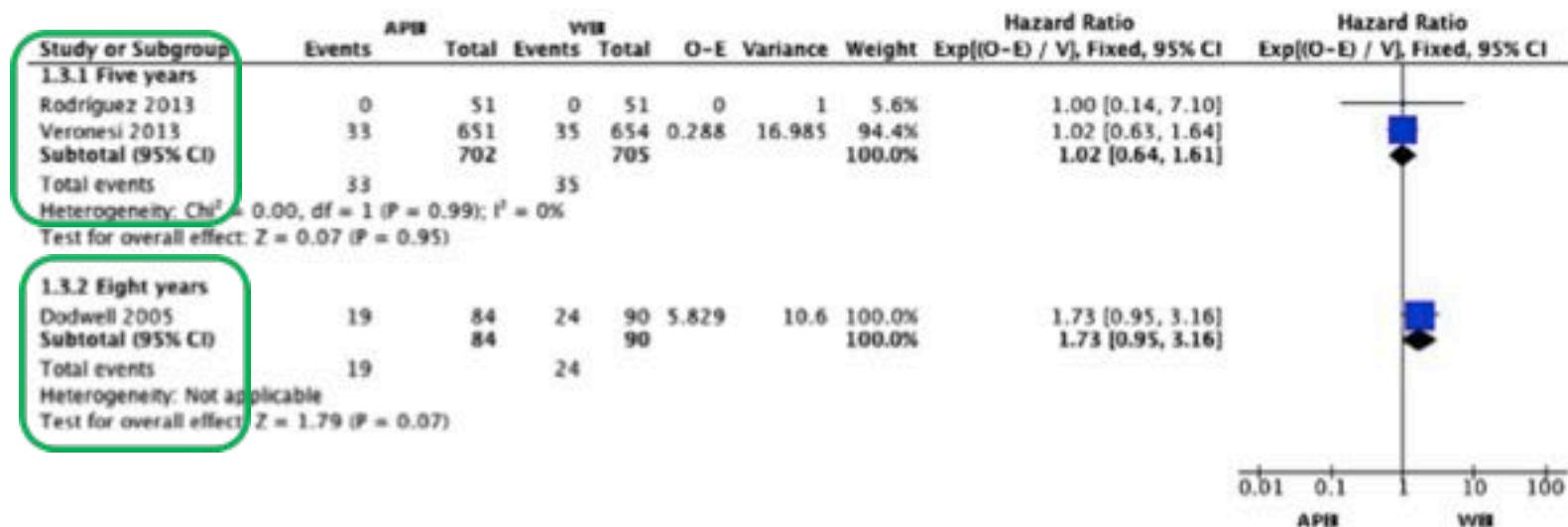


Fig. 3. Systemic recurrence.

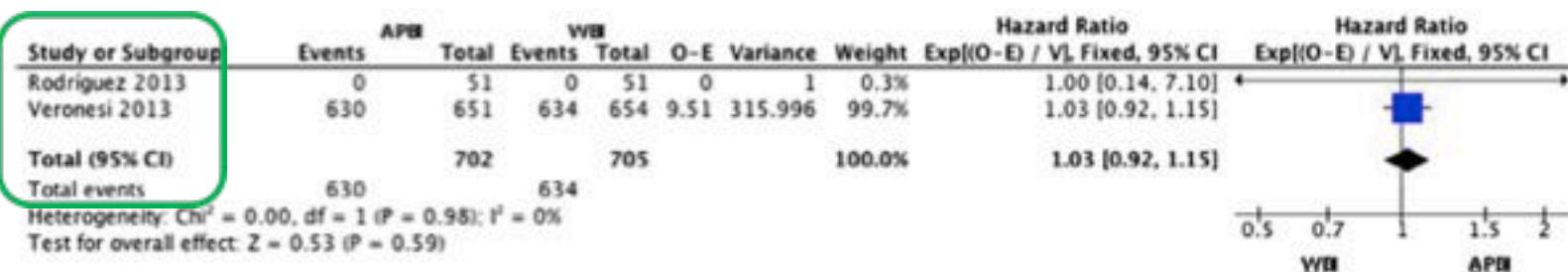


Fig. 4. Overall survival.

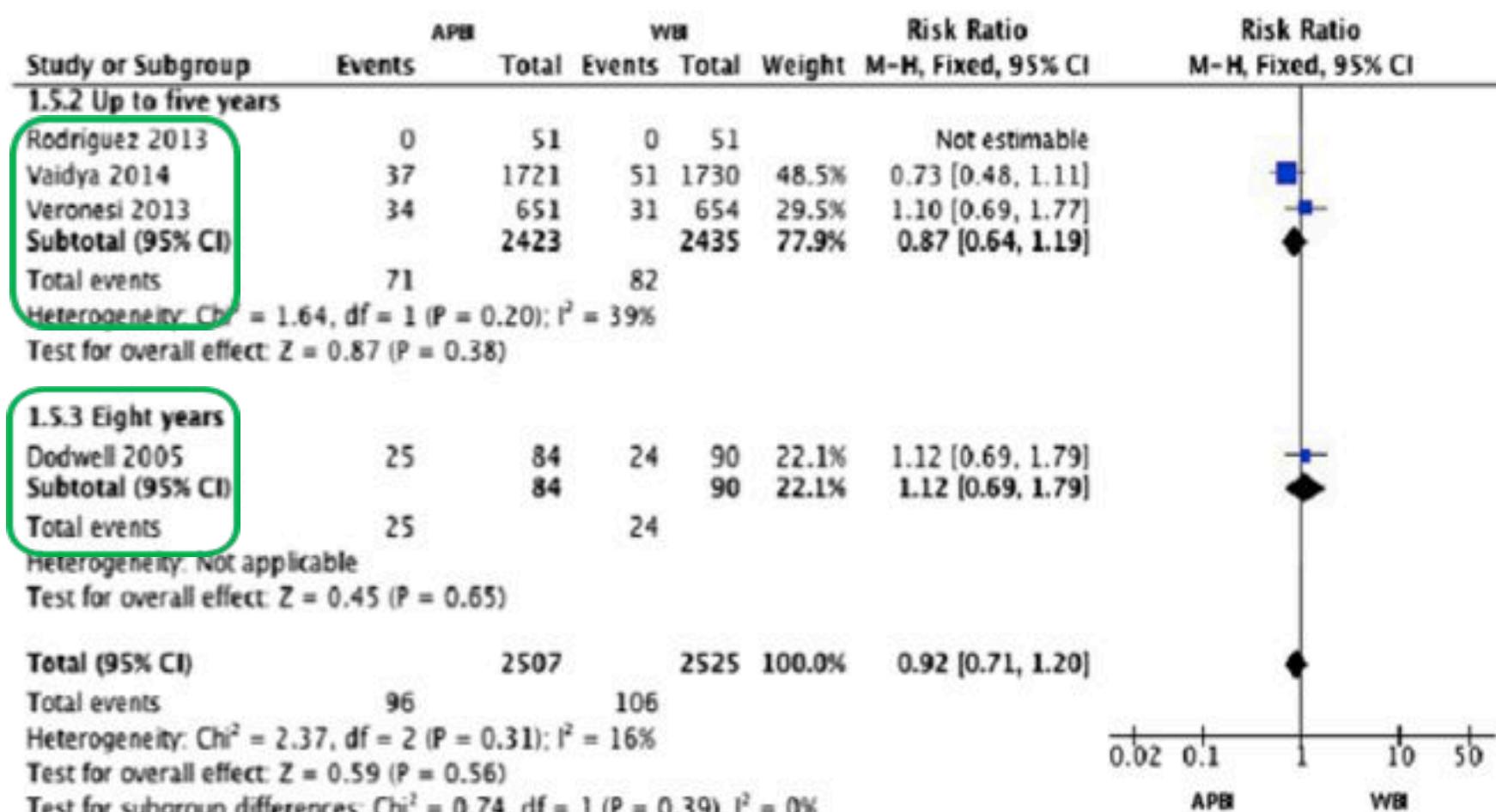


Fig. 5. Mortality.

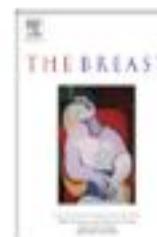
For Cosmesis, QoL, Toxicity: NS



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Original article

Accelerated partial breast irradiation using 3D conformal radiotherapy: Toxicity and cosmetic outcome



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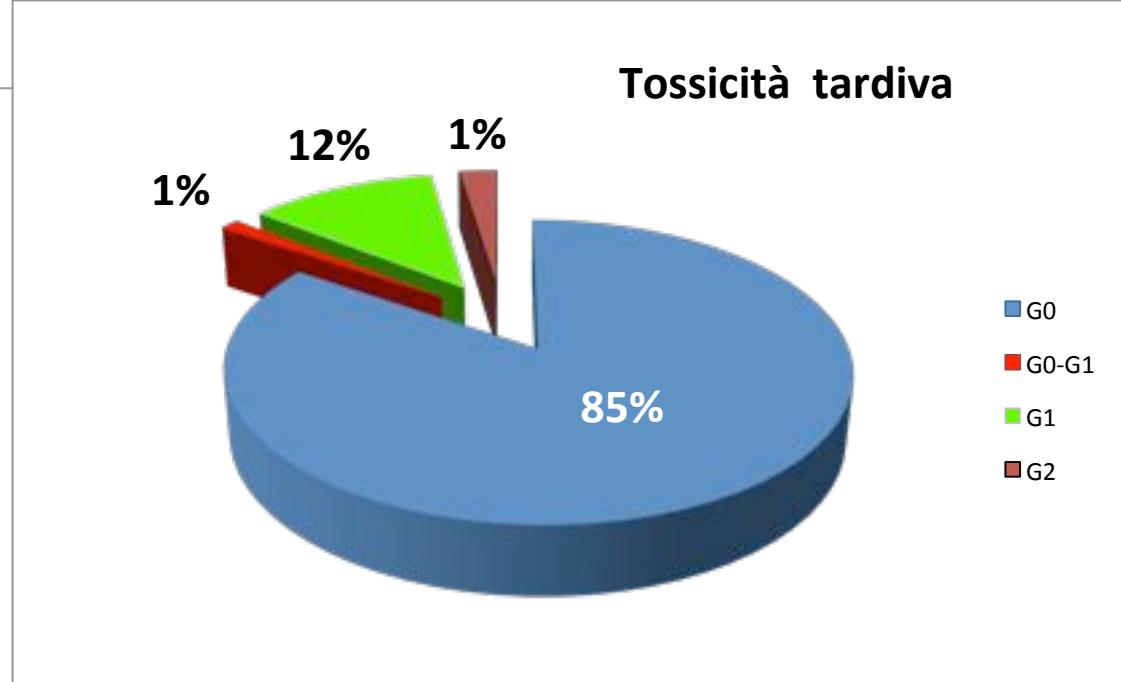
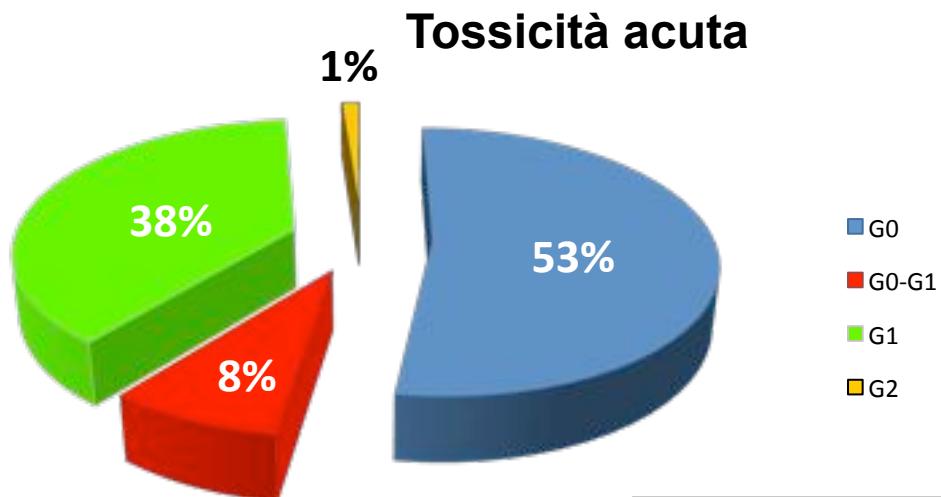
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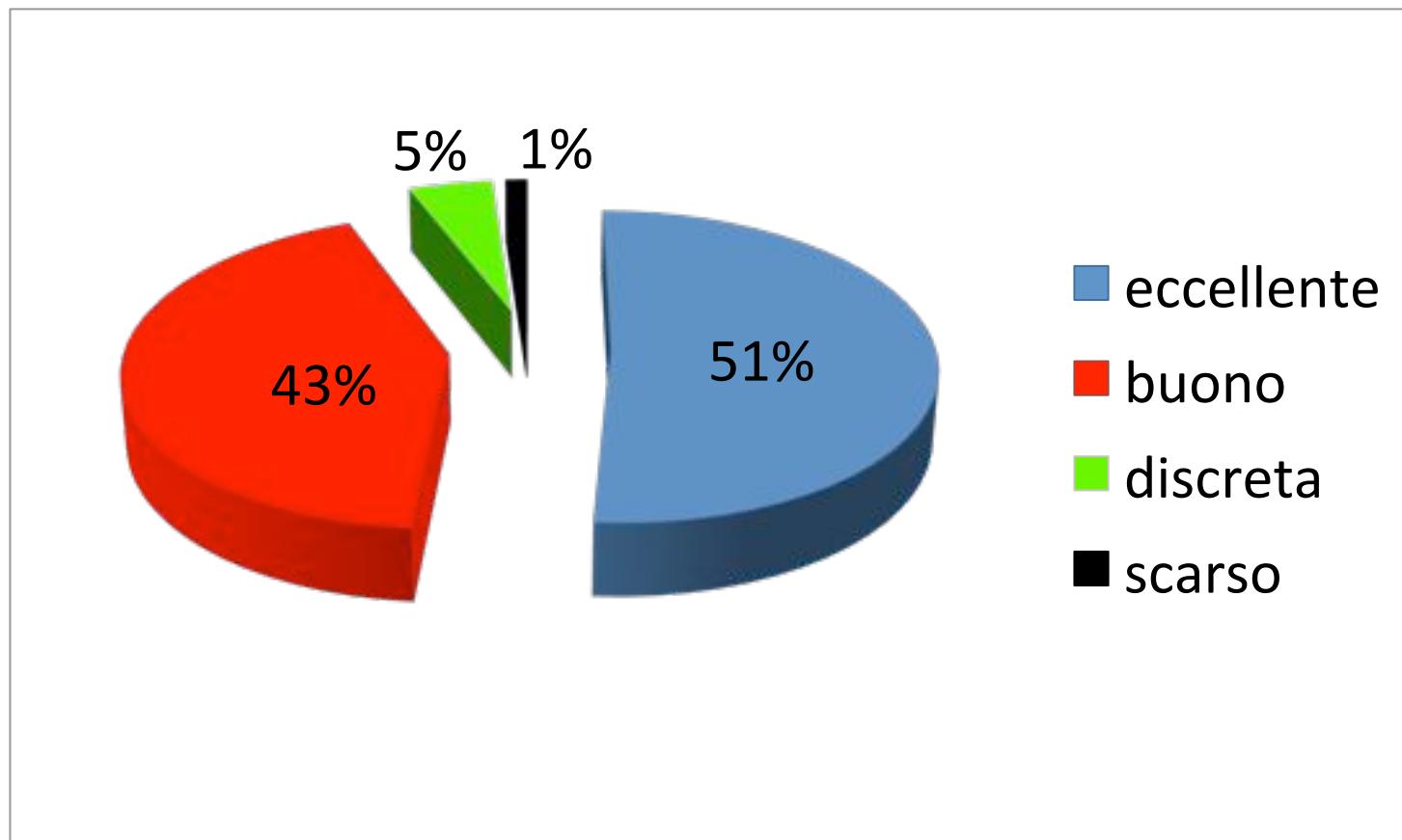
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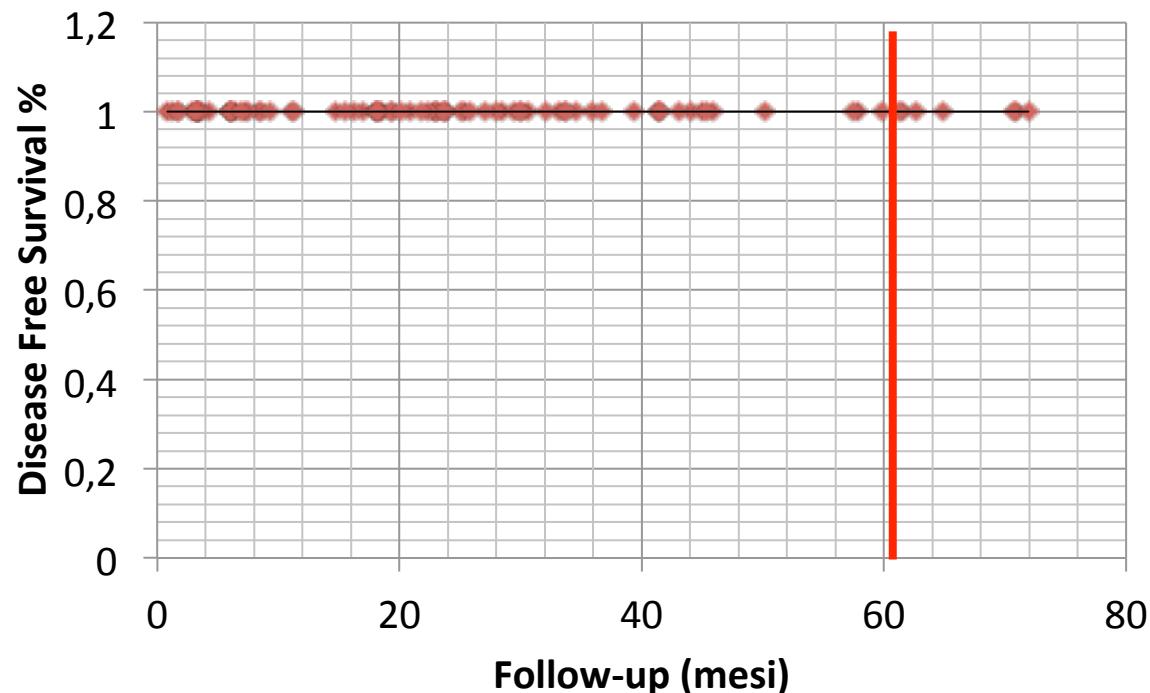
Tossicità cutanea acuta e tardiva



RESULTATO COSMETICO



Sopravvivenza libera da malattia



8% dei pazienti con controllo > 5 anni

CONCLUSIONI

- *L'APBI rispecchia la tendenza moderna a “trattare di meno” ma in maniera più precisa ed efficace aumentando l'indice terapeutico*
- *E' una tecnica che rispecchia il nuovo “credo” radioterapico: -volume + dose*
- *I volumi possono diminuire grazie ai progressi dell'imaging, alla conoscenza della malattia e allo screening*

CONCLUSIONI

- *L'aumento della dose è la condizione necessaria per l'eradicazione della malattia.*
- *L'individuazione ottimale del volume bersaglio consentita dalle nuove tecniche RT deve corrispondere ad un'efficacia superiore e ad una diminuzione della dose somministrata ai tessuti sani*
- *Malgrado il progresso tecnico la dose somministrata al cuore nella “whole breast” permane un problema importante*

CONCLUSIONI

- *Gli studi di APBI di fase II e i primi risultati degli studi di fase III evidenziano globalmente un basso tasso di recidive locali e una tossicità acuta e tardiva accettabile, tuttavia il FU è ancora limitato.*
- *I risultati della IORT possono generare perplessità, tuttavia è la tecnica che richiede la selezione più accurata.*
- *In termini di tossicità globale la radioterapia esterna segna il passo nei confronti della brachiterapia, dove però la selezione delle pazienti è più stretta:*
 -  *mammella grande, e tumori in prossimità della cute o della parete toracica.*

CONCLUSIONI

- *Nelle pazienti anziane con tumori della mammella a basso rischio, l'APBI potrebbe essere un giusto compromesso tra l'astensione dal trattamento radioterapico proposto da alcune linee guida, e un'irradiazione di 4/5/6 settimane.*

- *Le Linee Guida ASTRO, ESTRO etc sembrano appropriate nell'aver individuato le pazienti candidabili alla PBI al di fuori di studi clinici.*

CONCLUSIONI

- *Non è proponibile un atteggiamento minimalista verso tutte le pazienti ma non è nemmeno più proponibile il trattamento “whole breast” in tutti i casi → personalizzazione del trattamento*
- *L'analisi sui possibili vantaggi economici dell'APBI sono discussi e controversi in letteratura (diverse politiche di rimborso e tecniche utilizzate)*
- *Certamente in termini di logistica, q.d.v. delle pazienti e impegno dei “caregiver” costituiscono un vantaggio non solo teorico.*