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3° Taller Internacional Multidisciplinario de Cáncer de Mama & 1° Simposio de Cáncer U ∢ Ginecológico & 1° Taller de Planificación y Control de Calidad para Radiocirugía "De la práctica a las bases teóricas" Radioterapia post reconstrucción



I have no conflicts of interest Instituto Zunino Fundación Marie Curie



Breast reconstruction & RT: past or present challenge?

1. Introduction

2. RT after breast reconstruction

3. Breast reconstruction after RT

4. Sequestitecheurzetanino

5. New developments Adarte ing research

6. Conclusions



1. Introduction: why mastectomy?

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Who should not undergo breast conservation?

Risk factors:

- > focally incomplete margins: x2
- < 35 years: x 2 (yes, but: see next slide)</p>
- no radiotherapy: x 3-6

In larger cancers → PST to be considered. Oncoplastic procedures to be available.

Finally: remains an individual decision



Poortmans P, et al. Breast. 2017;31:295-302.

EORTC 10801 & DBCG-82TM.

Cumulative LRR according to histologic type:



Dutch population based cancer registry

2000-2004 cohort: 37,207 patients

A - 58.4% BCT logrank p<0.001 - 41.6% MRM 0.8 T1-2N0-1 Institu **UNINU** 0.0 2 10 12 14 8

Dutch population based cancer registry

2000-2004 cohort: 37,207 patients

Breast conserving therapy A - 58.4% BCT - 41.6% MRM Mastectomy 0.8 Cumulative overall survival T1-2N0-1 Institu Marie Curie MAST BCT 0.0 10 12 14 2 6 8

The Breast 35 (2017) 32-33



"Sometimes patients demand a mastectomy, driven by fear and the desire of getting rid of the disease while ignoring all this new information. It is important to inform them properly that, in most cases, breast cancer can be cured maybe even better without the need to be separated from of their breasts."

1. Introduction: why PMRT?

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Effect of radiotherapy after mastectomy and axillary surgery on 10-year recurrence and 20-year breast cancer mortality: meta-analysis of individual patient data for 8135 women in 22 randomised trials

EBCTCG (Early Breast Cancer Trialists' Collaborative Group)* Lancet 2014; 383: 2127-35 Fundación Marie Curie

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Figure 1.

Immediate Breast Reconstruction Rate from 2000–2010 in Patients Requiring Radiation. (6,875 reconstructions/40,568 women with BC)



Figure 2.

Reconstruction Rates by Method in Radiated Patients from 2000–2010 out of all specified reconstructions

USA-Low risk/BCS eligible

- Breast reconstruction increased from 11.6% to 36.4%
- Bilateral mastectomy for unilateral disease increased from

1.9% to 11.2% (not correlating with the rates of BRCA/PALB2) Fundación Marie Curie



USA-Low risk/BCS eligible

Breast reconst

Bilateral ma

1.9% to 11

Patients desire symmetry and cosmesis 6% to 36.4%

3RCA/PALB2)

increased from



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Modified from Ho A, et al. Lancet Oncol. 2017

Questions:

1. Is it safe (tumour control/survival)?

Is it safe (complications/cosmesis)?
 Is it feasible (RT issues)?
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Question 1:

Is it safe (tumour control/survival)?

Answer:

- Interval between surgery/chemotherapy & RT ~ acceptable
- Compromised DFS/OS??? More studies needed!
- Safe & well tolerated in LABC (Crisera Plast Reconstr Surg 2011;128:32-41)



Question 2:

Is it safe (complications/cosmesis)?

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Breast reconstruction & RT: *RT after reconstruction Capsular contracture following IBR with implant.*





Breast reconstruction & RT: *RT after reconstruction Capsular contracture following IBR with implant.*





Autologous tissue reconstruction vs implant.

	Implant	Autolog	Complic	Reoperation	Med FU
	No	No	%	%	mths
Philadelphia 2004		26	TRAM:0 Implant:5	2 (implants)	28
Boston 2002	Funda	ación I	TRAM:12 Implant:53	TRAM:8 Implant:42	32
Long Island 2008	69	23	ATR:9 Implant:55	ATR:0 Implant:19	38



Autologous tissue reconstruction vs implant.





Question 2:

Is it safe (complications/cosmesis)?

<u>Answer:</u>

- Increased capsular contraction (16% → 40%)
- Lower risk for complications after autologous IBR

Question 3:

• Is it technically feasible (RT issues)?

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Technical challenges:

- Chest wall shape
- Coverage of target volume
- Tissue heterogeneities
- Combination with regional nodes irradiation
- Avoidance of OAR (lung and heart)
- Delay in initiation of radiotherapy

Chest wall shape.





Coverage of target volume; regional RT.



Bilateral reconstruction.



Tissue heterogeneities.





Coverage/regional/OAR.

Impact of IBR (auto) on dosimetry (MDACC)

- N = 112
- Modified radical mastectomy
 → IBR (auto) (96% TRAM)
- N = 106 N =
- Modified radical mastectomy without IBR

➔ Dosimetric comparisons

Motwani SB. IJROBP 2006;66:76-82.
Coverage/regional/OAR.



New RT techniques.

<u>3D-CRT:</u>



New RT techniques.

<u>3D-CRT:</u>

-More experience

-No "low dose bath"



Volumetric IMRT:

- -Better TV coverage
- -Improved dose homogeneity



Breast reconstruction & RT: RT after reconstruction New RT techniques.











Question 3:

• Is it technically feasible (RT issues)?

Answer:

- Quite a challenge to meet all constraints
- More time and resource consuming
- Target volume delineation (!!!)



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Breast reconstruction & RT: Reconstruction after RT

Question:

• Is it safe (complications/cosmesis)?

Answers:

- Autologous in general preferred
- Individualise dación Marie Curie

Breast reconstruction & RT: past or present challenge?

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A Systematic Review of Complications of Implant-Based Breast Reconstruction with Pre-Reconstruction and Post-Reconstruction Radiation Therapy

Adeyiza O. Momoh, MD¹, Raouf Ahmed, BSc Hons², Brian P. Kelley, MD³, Oluseyi Aliu, MD³, Kelley M. Kidwell, PhD⁴, Jeffrey H. Kozlow, MD⁵, and Kevin C. Chung, MD MS⁶

<u>1006 → 26 articles</u> <u>1684</u> <u>patients/breasts</u> <u>ación</u> Marie Curie

- 14 on pre-reconstruction RT
- 23 on post-reconstruction RT

Reconstruction failures



Momoh AO, et al. Ann Surg Oncol. 2014;21:118–24

A Systematic Review of Morbidity Associated with Autologous Breast Reconstruction Before and After Exposure to Radiation Therapy- Are Current Practices Ideal?

Brian P. Kelley, MD¹, Raouf Ahmed, BSc Hons², Kelley M. Kidwell, PhD³, Jeffrey H. Kozlow, MD MS⁴, Kevin C. Chung, MD MS⁵, and Adeyiza O. Momoh, MD⁶

<u>897 \rightarrow 20 articles</u> 20 articles

- 6 on pre-reconstruction RT Marie Curie
- 9 on post-reconstruction RT
- 5 both

Wound healing complications



Kelley BP, et al. Ann Surg Oncol. 2014;21:1732–8

<u>= very similar:</u>

- But more complications than without PMRT
- Homologous better
- Carefully evaluate "quality" of (sub)cutaneous tissue Fundación Marie Curie

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Guidelines for PMRT- CTV_chest wall



Breast reconstruction & RT: *Developments* Guidelines for PMRT- CTV_chest wall after IBR

ESTRO ACROP consensus guideline for target volume delineation in the setting of

postmastectomy radiation therapy after implant-based immediate reconstruction for early stage breast cancer

Orit Kaidar-Person*, Birgitte Vrou Offersen*, Sandra Hol, Meritxell Arenas, Cynthia Aristei, Celine Bourgier, Maria Joao Cardoso, Boon Chua, Charlotte Coles, Tine Engberg Damsgaard, Dorota Gabrys, Reshma Jagsi, Rachel Jimenez, Anna M. Kirby, Carine Kirkove, Youlia Kirova, Vassilis Kouloulias, Tanja Marinko, Icro Meattini, Ingvil Mjaaland, Gustavo Nader Marta, Petra Witt Nystroem, Elzbieta Senkus, Tanja Skyttä, Tove F Tvedskov, Karolien Verhoeven, Philip Poortmans.



Breast reconstruction & RT: *Developments* Guidelines for PMRT- CTV_chest wall after IBR

- Target volume should be decided according to risk & site of recurrence.
- Understanding the surgical procedure and pathology report is essential!
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Breast reconstruction & RT: *Developments* Sites of recurrence

✓ 72-100% "chest wall" recurrences are Anterior Chest Wall Skin and subcutaneous tissue within the skin and subcutaneous tissue anterior to the pectoralis ²Harvard (n=17) = 100%-✓ #2nd most common site is within the Pectoralis Minor ³MDACC (n=60) = 93% pectoralis *Missouri (n=146) = 92% ✓ No reports of isolated rib/IC muscle Intercostals 75% recurrences intercostal muscles... 5Bryn Mawr (n=16)* Ribs Fundación Mar 72% Pectoralis Major 6MDACC (n=39)

Breast reconstruction & RT: *Developments* Sites of recurrence

- ✓ 72-100% "chest wall" recurrences are within the skin and subcutaneous tissue anterior to the pectoralis
- ✓ #2nd most common site is within the pectoralis
- No reports of isolated rib/IC muscle recurrences intercostal muscles...
- ... a few isolated rib and intercostal muscles are reported (Chang et al., Rad & Onc 2017)



Sites of recurrence – implant-based reconstruction



🕖 Vargo et al., IJROBP 2015



Vargo et al., IJROBP 2015

Sites of recurrence → target volumes

What are the target volumes in the immediate-reconstruction setting?

- Like without reconstruction but:
- Without the reconstructed (prosthesis, autologous (muscle, fat, skin), ...) material/tissue
- Some tissue might be displaced (pectoral muscles) or stretched (skin)

Sites of recurrence → target volumes

Yes, taking into account:

- Tissue that might be displaced (pectoral muscles)
- Tissue that can be stretched (skin)

Are the target volumes similar in case of implant-based reconstruction versus autologous flap?





Figure 1: CTV contouring of case with immediate breast reconstruction left using an implant. A: by writers of guideline of DBCG RT Recon Trial (n=5); B: by other radiation oncologists (n=18); C: by breast cancer surgeons (n=2).



Figure 2: Lymphatic draining pattern from the mammary region via the dermal plexus located within the subcutaneous tissues.



Figure 3: Implant positioning. A: retropectoral with full coverage by the pectoral muscle; B: retropectoral with partial coverage by the pectoral muscle and supportive material in the lower part; C: prepectoral with full coverage by supportive material.



Figure 4A: CTVp_chestwall with only a ventral part (red) in cases for whom only the subcutaneous lymphatic plexus should be irradiated. Pectoral muscles (yellow) and implant (green).



Figure 4B: CTVp_chestwall with a ventral (red) and dorsal (blue) part in cases for whom the subcutaneous lymphatic plexus should be irradiated as well as the part of the chest wall that was initially not covered by the pectoral muscles (yellow). Retropectoral implant (green).



Figure 4C: CTVp_chestwall with a ventral (red) and dorsal (blue) part in cases with a prepectoral implant (green). Pectoral muscles (yellow).



The DBCG RT Recon Trial:

Immediate versus delayed breast reconstruction in early breast cancer patients treated with mastectomy and adjuvant loco-regional radiation therapy.

A multicenter randomized clinical trial



Breast reconstruction & RT: Future research

Future research

- Target volume definition 1.
- In fact, this entire clinical question... 2.
- SSM/NSM +/- deflated tissue expander 3.
- (Free) fat grafting OZUNINO 4.
 - Fundación Marie Curie BRAVA & HBOT
- 5.
- RT before mastectomy 6.



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Breast reconstruction & RT: Conclusions

1. Advise BCT, unless...

- 2. Determine the odds for PMRT prior to surgery
- 3. No clear impact on prognosis
- 4. Impact on complications and cosmesis
- Less complications with autologous
 Individualise based on: RTeffects; BMI; HT; age; smoking;

stage; surgeondación Marie Curie

7. Joint multidisciplinary process & shared decision making



Breast reconstruction & RT: Conclusions

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