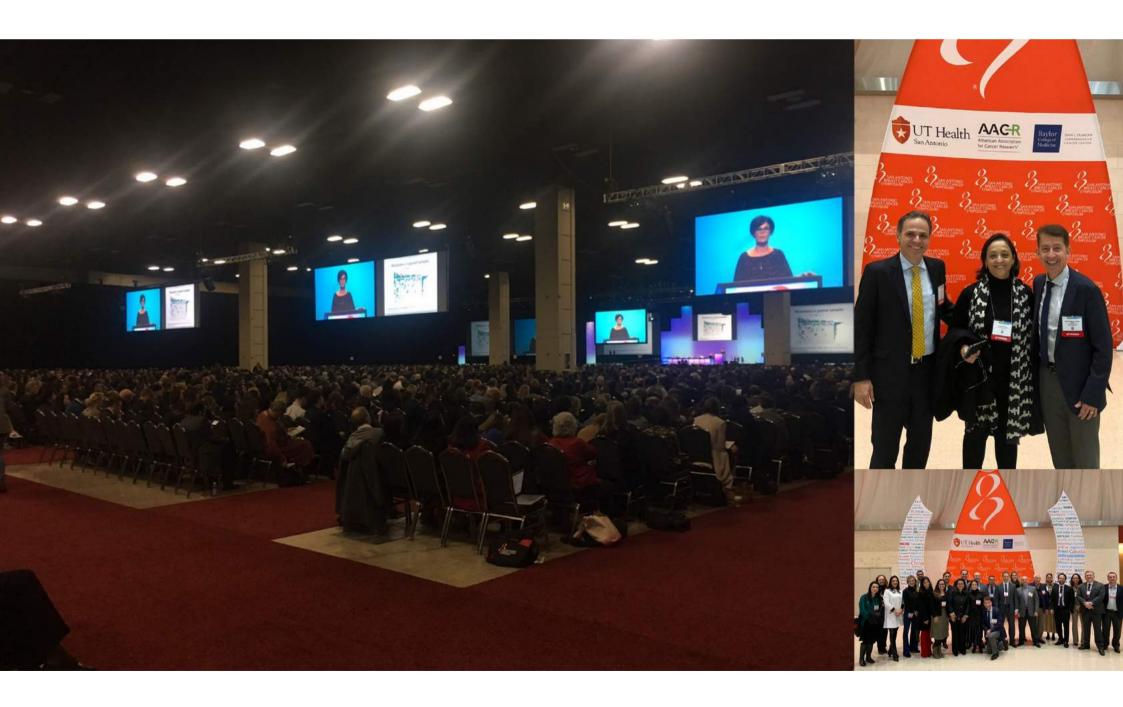
Novidades do SABCS 2018

Régis Resende Paulinelli, MD, PhD

Hospital das Clínicas - UFG

Hospital Araújo Jorge - ACCG



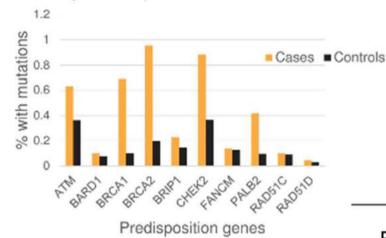


Age-related breast cancer risk estimates for the general population based on sequencing of cancer predisposition genes in 19,228 breast cancer patients and 20,211 matched unaffected controls from US based cohorts in the CARRIERS study

Fergus J. Couch, Ph.D. Mayo Clinic

San Antonio Breast Cancer Symposium - December 4-8, 2018

Frequency of mutations for known breast cancer predisposition genes (all races and ethnicities)

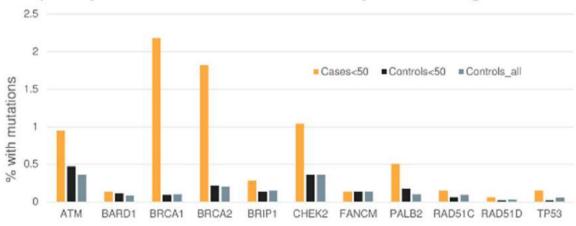


Case mutation frequency 4.2% Control mutation frequency 1.6%

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Frequency of mutations in cases =<50 years at diagnosis



Predisposition Genes

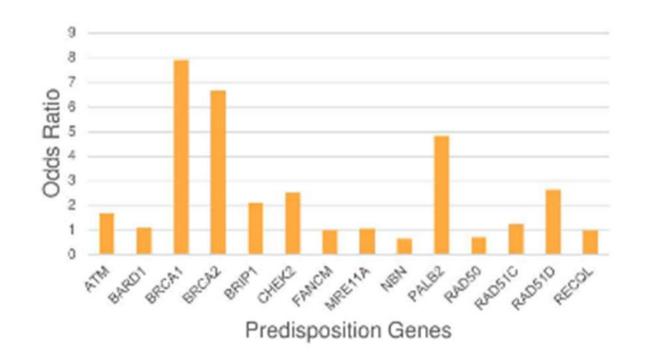
Case mutation frequency 7.3% Control mutation frequency 1.8%

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CARRIERS breast cancer risk estimates by panel gene

	Relative Risk	p-value
ATM	1.7	0.001
BARD1	1.1	0.80
BRCA1	7.9	< 0.001
BRCA2	6.7	< 0.001
BRIP1	2.1	0.01
CHEK2 (truncating)	2.5	< 0.001
FANCM	1.0	0.95
MRE11A	1.0	0.90
NBN	0.6	0.16
PALB2	4.8	< 0.001
RADSO	0.7	0.15
RADS1C	1.2	0.58
RADS1D	2.6	0.15
RECQL	1.0	0.89



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Radiotherapy or surgery of the axilla after a positive sentinel node in breast cancer patients: 10-year results of the EORTC AMAROS trial

By the EORTC Breast Cancer Group and Radiation Oncology Group In collaboration with the Dutch BOOG Group and ALMANAC Trialists' Group

Emiel J Rutgers

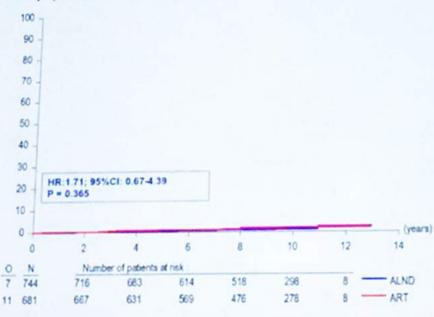
The Netherlands Cancer Institute, Amsterdam

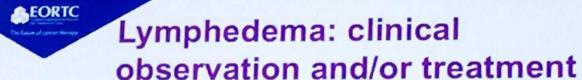


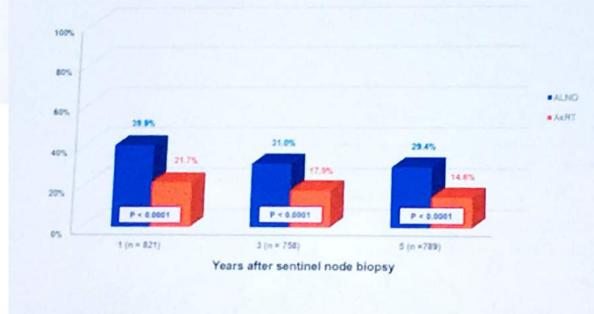


Axillary recurrence rate









F-value from exact Fisher's test-



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Regional node irradiation: Meta-analysis of 13,500 women in 14 trials

Early Breast Cancer Trialists' Collaborative Group (EBCTCG)

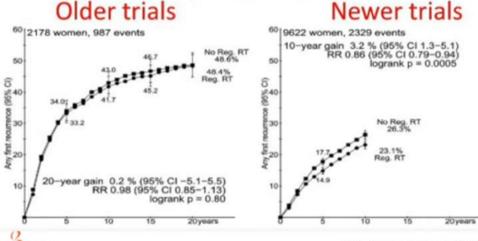
Writing Committee: David Dodwell (presenter), Carolyn Taylor, Paul McGale, Charlotte Coles, Fran Duane, Richard Gray, Thorsten Kühn, Christophe Hennequin, Robert Hills, Sileida Oliveros, Yaochen Wang, Jonas Bergh, Kathy Pritchard, Sandra Swain, Jens Overgaard, Philip Poortmans, Tim Whelan

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Any recurrence

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Older trials



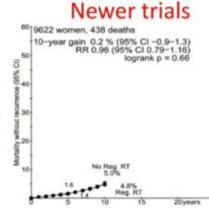
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Non-breast-cancer mortality

Older trials





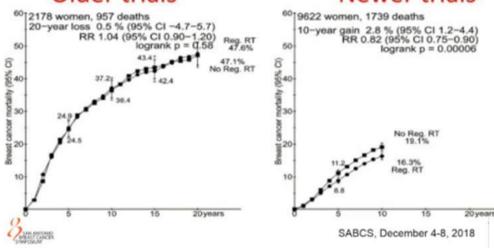


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Breast cancer mortality

Older trials Newer trials

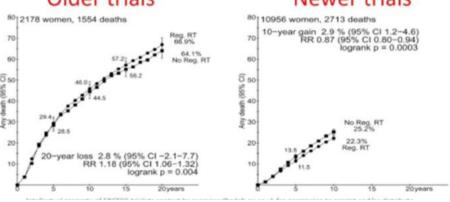


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Overall mortality



Newer trials





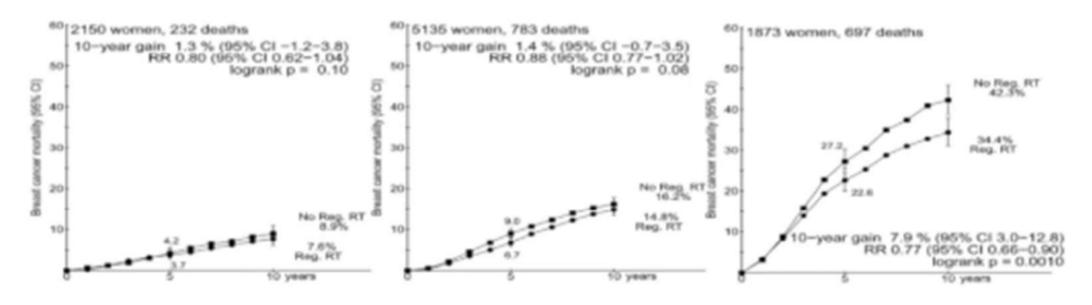
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Newer trials: Breast cancer mortality

pN0

pN1-3

5N4+





RAPID



Randomized Trial of Accelerated Partial Breast Irradiation using 3-Dimensional Conformal Radiotherapy (3D-CRT)

T Whelan, J Julian, M Levine, T Berrang, DH Kim, CS Gu, I Germain, A Nichol, M Akra, S Lavertu, F Germain, A Fyles, T Trotter, F Perera, S Balkwill, S Chafe, T McGowan, T Muanza, W Beckham, B Chua, I Olivotto, for the RAPID Trial Investigators

Ontario Clinical Oncology Group

SABCS, December 4-8, 2018







Primary results of NSABP B-39/ RTOG 0413 (NRG Oncology): A randomized phase III study of conventional whole breast irradiation (WBI) versus partial breast irradiation (PBI) for women with stage 0, I, or II breast cancer

F Vicini (NSABP PI), R Cecchini, J White (RTOG PI), T Julian, D Arthur, R Rabinovitch, R Kuske, D Parda, P Ganz, M Scheier, K Winter, S Paik, H Kuerer, L Vallow, L Pierce, E Mamounas, J Costantino, H Bear, I Germain, G Gustafson, L Grossheim, L Petersen, R Hudes, W Curran, N Wolmark

RAPID (N = 2,135, 18% DCIS)

- APBI 90% 3D conformal photons;
 10% IMRT
- WBI 16 fraction regimen, 20%
 Boost
- 100% node negative
- Median age 61 years old

- APBI 71% 3D conformal photons;
 29% brachytherapy
- WBI 25-28 fraction regimens, 80% boost
- 10% node positive
- Median age 54 years old



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- 10% node positive
- Median age 54 years old



APBI (1 week, BID) Main Findings

RAPID (N = 2,135)

- 8-yr IBTR WBI 2.8% vs APBI 3%,
 P= NS
- Non-inferiority endpoint met

NRG B39/0413 (N = 4,216)

- 10-yr IBTR WBI 4.1% vs APBI 4.8%,
 P = NS
- Non-inferiority endpoint <u>not met</u>
- Recurrence Free Interval 1.6% worse with ABPI, P = 0.02



IBTR by **PBI** Method

Treatment Group	# of Pts	# of Events	Hazard Ratio (HR)	HR 95% Confidential Interval	10-yr Cum Incidence
WBI	2,011	67	REF		3.8%
PBI					
Multi-catheter brachytherapy	130	9	2.21	1.10 - 4.46	7.7%
Single-entry brachytherapy device	358	24	2.15	1.34 - 3.44	7.8%
3DCRT (external beam)	1,535	55	1.04	0.73 - 1.49	3.7%

This analysis used a per-protocol population, which excluded those who did not receive their randomly assigned treatment



APBI (1 week, BID) Main Findings

- RAPID (N = 2,135)
 - 3-yr Grade 3 Toxicity WBI 1% vs APBI 4.5%, P < 0.001
 - · 7-yr fair/poor cosmesis more common in the APBI arm



San Antonio Breast Cancer Symposium, December 4-8, 2018









in partnership with

Dose escalated simultaneous integrated boost radiotherapy for early breast cancer: 3-year adverse effects - IMPORT HIGH trial (CRUK/06/003)

Dr Charlotte Coles, Clare Griffin, Anna Kirby, Joanne Haviland, Jenny Titley, Kim Benstead, Adrian Murray Brunt, Charlie Chan, Laura Ciurlionis, Omar Din, Ellen Donovan, David Eaton, Adrian Harnett, Penelope Hopwood, Monica Jefford, Peter Jenkins, Caroline Lee, Mary McCormack, Liz Sherwin, Isabel Syndikus, Yat Tsang, Nicola Twyman, Ramachandran Ventikaraman, Sairanne Wickers, Maggie Wilcox, Judith Bliss and John Yarnold

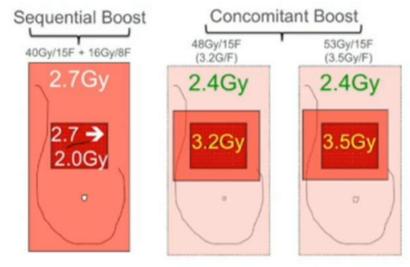
On behalf of the IMPORT HIGH Trial Management Group

San Antonio Breast Cancer Symposium Thursday 6th December, 2018



San Arkonio Breast Cancer Symposium, December 4-8, 2018

TRIAL DESIGN: Dose Escalated Intensity Modulated RT



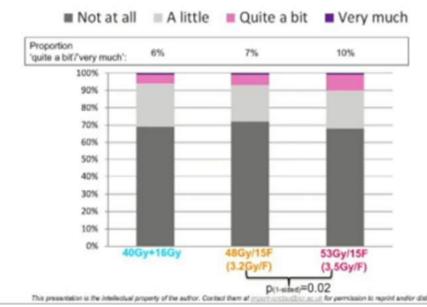
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San Antonio Breast Concer Symposium, December 4-8, 2018

ENDPOINTS: CRO: breast induration at 3 years





San Antonio Breast Cancer Symposium*, December 4-8, 2018

The impact of breast cancer surgery on quality of life: Long term results from E5103

Shoshana M. Rosenberg, Anne O'Neill, Karen Sepucha, Kathy D. Miller, Chau T. Dang, Donald W. Northfelt, George W. Sledge, Bryan P. Schneider, Ann H. Partridge

ECOG – ACRIN E5103 Bevacizumabe 4.994 pacientes.





San Antonio Breast Cancer Symposium*, December 4-8, 2018

QoL analysis: BCS vs. Mastectomy

	Mear	P**	
	BCS	Mastectomy	
FACT-B	114	109	0.01
EQ-5D-3L Index	0.84	0.80	0.04
EQ-VAS	82	78	<0.01

Minimally important differences:

FACT-B: 7-8 points

*Higher scores=Better QoL

EQ-5D-3L Index: 0.06 points

**Wilcoxon rank sum test p-value EQ-VAS: 7 points



San Antonio Breast Cancer Symposium*, December 4 -8, 2018

Local therapy and quality of life outcomes in young women with breast cancer

Laura Dominici, Jiani Hu, Tari King, Kathryn J. Ruddy, Rulla M. Tamimi, Jeffrey Peppercorn, Lidia Schapira, Virginia F. Borges, Steven E. Come, Ellen Warner, Ann Partridge, Shoshana Rosenberg

The Young Women's Breast Cancer Study (YWS)

- · Multicenter, prospective cohort study
- 12 participating hospitals (academic and community)





BREAST-Q Mean Scores

100

80

60

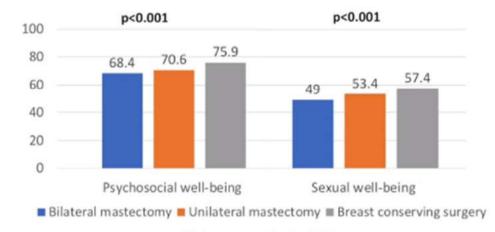
40

20

0

p= 0.008 p= 0.8 78.7 78.9 78.9 60.4 59.3 65.5

BREAST-Q Mean Scores



Higher score = Better QOL

Satisfaction with breasts

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■ Bilateral mastectomy ■ Unilateral mastectomy ■ Breast conserving surgery

Higher score = Better QOL

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Multivariate Analysis: Satisfaction with Breasts

	β*(95% CI)	P-value**
Surgery factors:		
Unilateral mastectomy vs. BCS	-8.7 (-13.1, -4.3)	<0.001
Bilateral mastectomy vs. BCS	-9.3 (-14.4, -4.2)	<0.001
Other treatment factors:		
Radiation: Yes vs. No	-7.5 (-11.3, -3.6)	<0.001
Patient factors:		
Financial status: Uncomfortable vs. comfortable	-5.4 (-9.8, -1.0)	0.02

^{*}B Difference in BREAST-Q score

Physical well-being

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^{**}Not significant for age, race, marital status, work status, education level, stage, chemotherapy, endocrine therapy, time since surgery, lymphedema



Surgical treatment after neoadjuvant systemic therapy in young women with breast cancer: Results from a prospective cohort study

Hee Jeong Kim ^{1,2}, Laura Dominici^{1,3}, Shoshana Rosenberg¹, Linda Ma Pak^{1,3}, Phillip D. Poorvu¹, Kathryn Ruddy¹, Rulla Tamimi³, Lidia Schapira⁵, Steven Come⁵, Jeffrey Peppercorn⁷, Virginia Borges³, Ellen Warner², Hilde Vardeh⁵, Laura Collins⁵, Rachel Gaither¹, Tari King^{1,3}, Ann H. Partridge¹

¹Dana-Farber Cancer Institute, Boston, MA; ²Asan Medical Center, Seoul, South Korea; ⁴Brigham and Women's Hospital, Boston, MA; ⁴Mayo Clinic, Rochester, MN; ²Stanford University, Palo Alto, CA; ⁵Beth Israel Deaconess Medical Center, Boston, MA; ⁵ Massachusetts General Hospital, Boston, MA; ⁸University of Colorado Cancer Center, Aurora, CO; ⁹ Sunnybrook Health Science center, Toronto, ONT





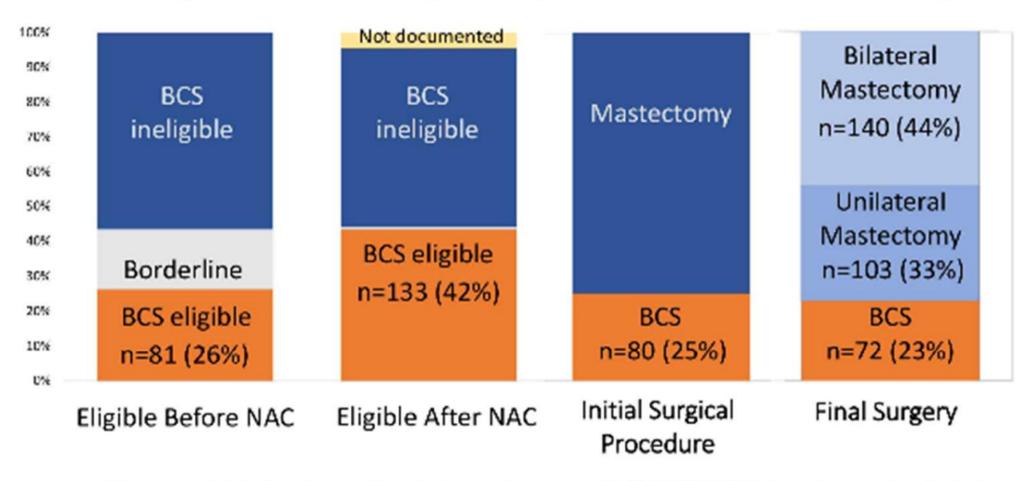


The Young Women's Breast Cancer Study (YWS)

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Change in BCS eligibility after NAC and surgery





Reasons for choosing mastectomy in BCS-eligible patients (N=55)

Unknown n=4 (7%) Family history n=3 (5%)

Mutation+

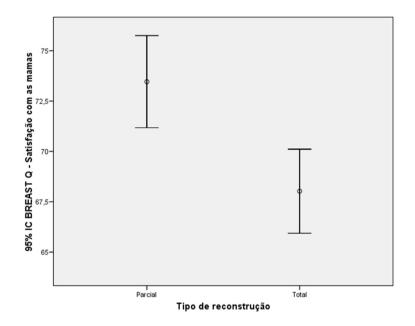
n=19 (35%)

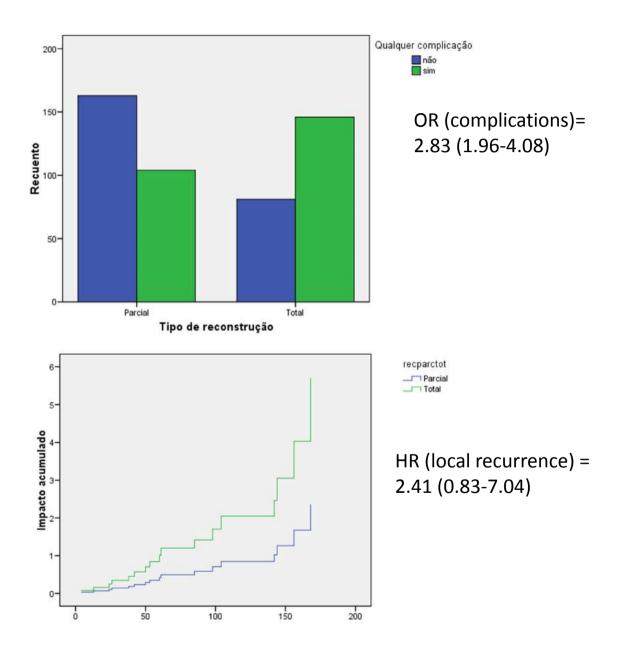
Preference* n=29 (53%)

- The most common documented reason that BCS-eligible patients chose mastectomy was patient preference (53%)
- 40% chose mastectomy because of carrying a BRCA 1 or 2, or p53 mutation or having a strong family history
- 75% who chose mastectomy underwent bilateral mastectomy
- Among BCS-eligible patients with cCR and/or ultimately pCR who chose mastectomy, these reasons were similar

^{*}Preference was defined as someone who chose mastectomy without having a mutation or strong family history

• 509 pacientes do consultório/hc/haj





PD8-01 Microscopic extracapsular extension in sentinel lymph nodes does not mandate axillary dissection in Z0011-eligible patients

Barrio AV, Downs-Canner S, Cody HS, Van Zee KJ, Gemignani ML, Pilewskie M, Plitas G, El-Tamer M, Kirstein L, Capko D, Patil S, Morrow M. Memorial Sloan Kettering Cancer Center, New York, NY.

Background

In ACOSOG Z0011 and AMAROS, matted nodes with gross extracapsular extension (ECE)—a risk factor for locoregional recurrence—were an indication for axillary dissection (ALND), but the effect of *microscopic ECE* (mECE) in the sentinel nodes (SLNs) on recurrence was not examined.

Methods

Between 2010-2017, 815 patients with cT1-2N0 breast cancer and SLN metastasis were prospectively managed according to Z0011 criteria, with ALND for those with >2 positive SLNs. Management of mECE was not specified. Here we report outcomes of patients with 1-2 positive SLNs treated with SLN biopsy alone (n=685) and evaluate the impact of mECE on nodal recurrence. Outcomes of the 118 patients treated with ALND, of which 70% had >2 positive SLNs, are provided for comparison.

Results

Median patient age was 58 years and median tumor size was 1.7 cm. In the SLN group, 210 (31%) had mECE. Patients with mECE were older, had larger tumors, were more likely to be hormone receptor positive (HR+) and HER2-, have 2 positive SLNs, and to receive nodal radiation. At a median follow-up of 41 months, no isolated axillary failures were observed. There were 11 nodal recurrences; 2 isolated, 4 synchronous with breast, and 5 with distant failure. The 5-year rate of any nodal recurrence was 1.6% and did not differ by mECE (2.3% vs 1.3%, p=0.84). No differences were observed in local (0% mECE vs. 1.9% no mECE, p=0.08) or distant (1.2% mECE vs. 4.6% no mECE, p=0.31) recurrence rates by mECE status. In comparison, in the 118 patients having ALND, 101 (86%) had mECE, and 1 combined nodal and distant recurrence was seen.

Conclusions

In Z0011-eligible patients, rates of nodal recurrence in patients with mECE are low after treatment with SLN biopsy alone, even in the absence of routine nodal radiation. The presence of mECE should not be considered a routine indication for ALND.

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Organização e supervisão

Dr. Maurício Resende (SE) Dr. Cícero Urban (PR) Dr. Regis Paulinelli (GO) Dias **30/11 e 1°/12**, em **Curitiba (PR)**

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