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San Francisco, CA  
United States  
January 27, 2018



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San Francisco, CA  
USA  
January 27, 2018

# Advances in Approaches to Local-Regional Disease



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Disclosure(s)

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Research Support:	N/A
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Advisory Panel/Consultant:	Genomic Heath
Co-founder/Stockholder:	N/A
Employee (part time):	N/A

# Best Of SABCS 2017

## Local-regional Treatment

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# Impact of the SSO-ASTRO Consensus Guidelines on Invasive Margins on the Re-excision Rate among Patients Undergoing BCS

Maria Chavez-MacGregor, et al, MD Anderson

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### Impact of the SSO-ASTRO Consensus Guidelines on Invasive Margins on the re-excision rate among patients undergoing breast conserving surgery (BCS).

Mariana Chavez-MacGregor<sup>1,2</sup>, Xiudong Lei<sup>1</sup>, Monica Morrow<sup>3</sup> and Sharon H. Giordano<sup>1,2</sup><sup>1</sup>Department of Health Services Research, The University of Texas MD Anderson Cancer Center, <sup>2</sup>Department of Breast Medical Oncology, The University of Texas MD Anderson Cancer Center, <sup>3</sup>Department of Surgery, Memorial Sloan Kettering Cancer Center.

## Background

- BCS has been historically associated with a high re-excision rate, driven in part by lack of consensus on what constitutes an adequate negative margin.
- The SSO-ASTRO consensus guideline on Invasive margins defined a negative margin as no ink on tumor.
- Evidence suggests that more widely clear margins do not further decrease the risk of recurrence.
- The SSO-ASTRO consensus guideline on Invasive margins can potentially reduce the need for re-excision.
- In a large nationwide cohort of breast cancer patients undergoing BCS for Invasive breast cancer we evaluate the rates of re-excision following BCS before and after the SSO-ASTRO consensus guidelines were disseminated.

## Methods

- Breast cancer patients undergoing BCS for invasive breast cancer between January 2012 and December 2015 were identified among female beneficiaries in the MarketScan database.
- Patients receiving chemotherapy before surgery were excluded.
- Based upon presentation of the guideline recommendations in October 2013, the pre-guideline period was defined from January 2012 to September 2013. On-line publication of the guideline in February 2014 led to definition of the post-guideline period from March 2014 onwards. The peri-guideline period was defined as the time between the pre and post-guideline intervals.
- Any re-excision or mastectomy within 3 months of initial BCS was identified using ICD-9 or CPT codes.
- Overall re-excision rates and 95% CI were calculated; groups were compared using X<sup>2</sup> test.
- We used a regression model to evaluate the association between pre-peri-post guideline period and re-excision while adjusting for important covariates.
- Results are expressed as risk ratios (RRs) and 95%CI.

## Results

- 38,573 patients were included. The overall re-excision rate was 23.9%. The pre-guideline re-excision rate was 25.3% compared to 21.6% in the post-guideline period.
- The rate of mastectomy as the final surgical procedure was 20.2% in the pre-guideline period and 19.1% in the post-guideline ( $p=0.15$ ).

TABLE-1. Comparison of overall re-excision rates stratified by study period

	Patients	Re-excitation events	Mean (95% CI)	P
Pre-guideline (Jan 2012 to Sept 2013)	25,138	5111	0.203 (0.197, 0.209)	<0.0001
Post-guideline (October 2013 to February 2014)	4857	1113	0.228 (0.226, 0.234)	
Post-guideline (March 2014 to Sept 2015)	13,827	2980	0.216 (0.213, 0.220)	
Coverall	38,513	9054	0.230 (0.226, 0.234)	

- We observed significant geographic variability by state in the decrease of the re-excision rates.
- Patients undergoing BCS in the post-guideline period had a statistically significant decrease in the risk of re-excision compared to patients undergoing surgery in the pre-guideline period (RR=0.87; 95%CI 0.84-0.91; p<0.001).

FIGURE-1. Quarterly re-excitation rates

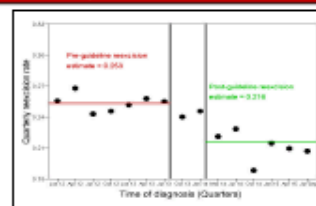


TABLE-2. Patient and clinical characteristics according to study period.

	Pre-pandemic (n = 2016)	Post-pandemic (n = 402)	Pre-pandemic (n = 1007)	p	
<b>Demographics</b>					
Sex					
Male	1024	74.8	2464	75.8	
Female	911	28.4	1115	24.2	<0.001
Age category					
18-24	36.9	18.6	88	16.6	0.078
25-34	217.6	10.7	462	10.9	0.277
35-44	270.3	16.8	666	27.6	0.001
45-54	307.1	16.2	856	16.6	0.260
55-64	112.3	4.0	247	5.2	0.001
65-74	269.1	13.5	656	14.1	0.111
≥ 75	88	4.4	176	3.3	0.277
<b>Economic status</b>					
Income					
< 1142.2	71.8	33.68	71.8	34.8	0.000
≥ 1142.2	304.8	8.48	304.8	30.0	0.177
Yes	198	7.4	365	7.7	0.162
<b>Employment status</b>					
Employed	917	82.8	2446	81.1	0.075
Unemployed	249.7	17.1	762	16.8	0.264
<b>Health status</b>					
Positive	107.1	62.8	364	61.1	0.016
Negative	249.7	17.1	762	16.8	0.264
<b>Exercise frequency</b>					
Frequency					
Yes	839	62.2	2262	64.4	0.141
No	339	26.8	1286	27.7	0.024
Yes	1038	21.4	1028	22.1	0.880
<b>Exercise intensity</b>					
Intensity					
Low	698	94.2	4315	89.6	0.000
High	110	6.8	261	6.4	0.177
<b>Exercise type</b>					
Frequency					
PAC	112.8	58	2656	67.2	0.001
Other	102.2	13.2	548	12.6	0.001
Intensity					
Low	876.2	27.8	1987	26.7	0.470
Other	67.8	4.3	168	4.1	0.311
<b>Region</b>					
Region					
North-west	42.9	26.8	1091	23.4	0.028
North-east	66	26	1124	26.1	0.028
South	816	26.8	1452	31.4	0.001
South-east	24.9	22.2	188	26.1	0.001
Unknow	8.8	2.2	12	2.6	0.171

POLUPB-2. Decrease in the re-excitation rate by state



TABLE-3. Multivariable model evaluating the determinants of re-escalation

	Relative Mean	95% CI	P
Preparation			
Fast preparation	1.00	0.50-1.01	0.1
Slow preparation	0.87	0.30-0.91	<0.001
Age at surgery			
18-49	1		
50-59	0.84	0.60-1.1	0.347
60-69	0.9	0.69-1.08	0.301
70-79	0.84	0.58-1.06	0.012
80-89	0.61	0.37-0.91	<0.001
≥90	0.62	0.37-1.06	<0.001
N	0.68	0.59-0.77	<0.001
Dependability			
1	0.86	0.70-1.0	0.78
2	1.08	0.96-1.12	0.2
English words score			
Highly	1.27	1.21-1.32	<0.001
Positive			
Dependability	1.27	1.21-1.32	<0.001
Yes			
Yes	0.8	0.77-0.84	<0.001
No	1.08	0.96-1.12	0.26
Dependence			
FPG	1.03	0.97-1.07	0.363
HMO	0.81	0.68-0.97	0.002
Yes	0.84	0.69-1.0	0.002
Region			
Northwest	1.03	0.97-1.06	0.3
Southwest	1.03	1.07-1.07	<0.001
Other	1.08	1.06-1.10	<0.001

## Conclusions

- There has been a statistically significant decrease in the re-excision rate after BCS associated with the dissemination of the SSO-ASTRO consensus guideline on invasive margins.
- Our study confirms the impact that guidelines have modifying patterns or practice, reducing the frequency of unnecessary interventions.
- Our study evaluates early implementation; likely underestimating the ultimate impact of the guideline.
- In 2014 there were approximately 113,894 BCS nationwide. A decrease in the re-excision rate of 3.7% corresponds to 4,214 fewer unnecessary procedures just in one year. The comorbidity and the cost associated with this reduction has profound implications for patients and for the health care system.

# Methods

- Breast cancer pts undergoing BCS for invasive disease between Jan 2012 and Dec 2015 identified among beneficiaries in MarketScan database
- Based on guideline recommendations in October 2013 (presentation)
  - Pre-guideline period Jan 2012-Sept 2013
  - Post-guideline period March 2014 onward (publication Feb 2014)
- Re-excision or completion mastectomy within 3 months of initial surgery using CPT codes and ICD-9 codes

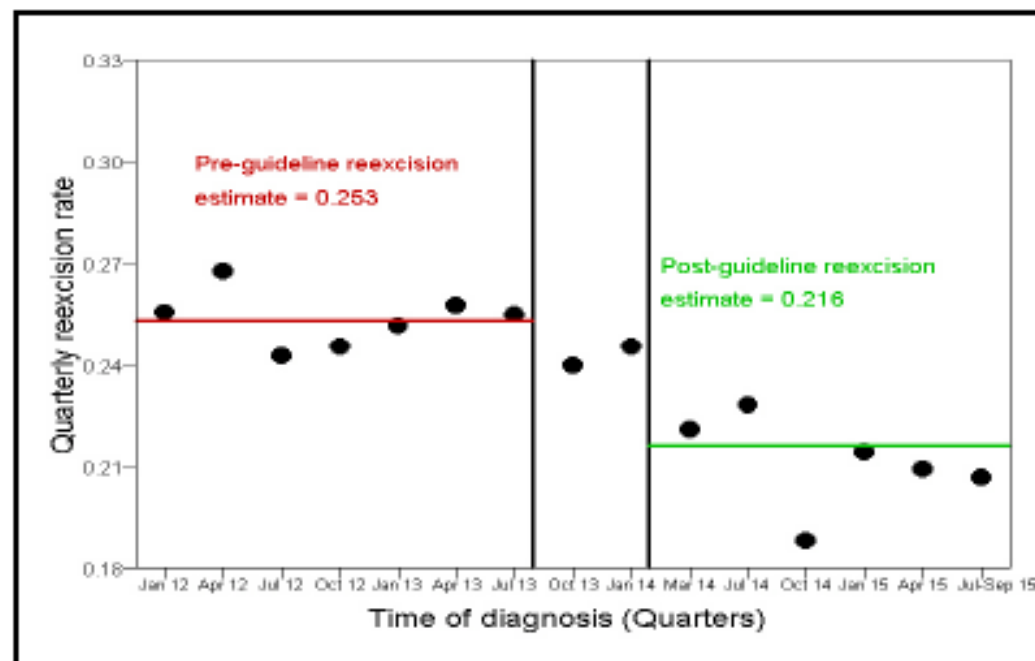
# Results

**TABLE-1. Comparison of overall re-excision rates stratified by study period**

	Patients	Re-excision events	Mean (95% CI)	P
Pre-guideline (Jan 2012 to Sept 2013)	20,159	5111	0.253 (0.247,0.259)	<0.0001
Peri-guideline (October 2013 to February 2014)	4607	1113	0.242 (0.229, 0.254)	
Post-guideline (March 2014 to Sept 2015)	13,807	2980	0.216 (0.209,0.223)	
Overall	38,573	9204	0.239 (0.234, 0.243)	

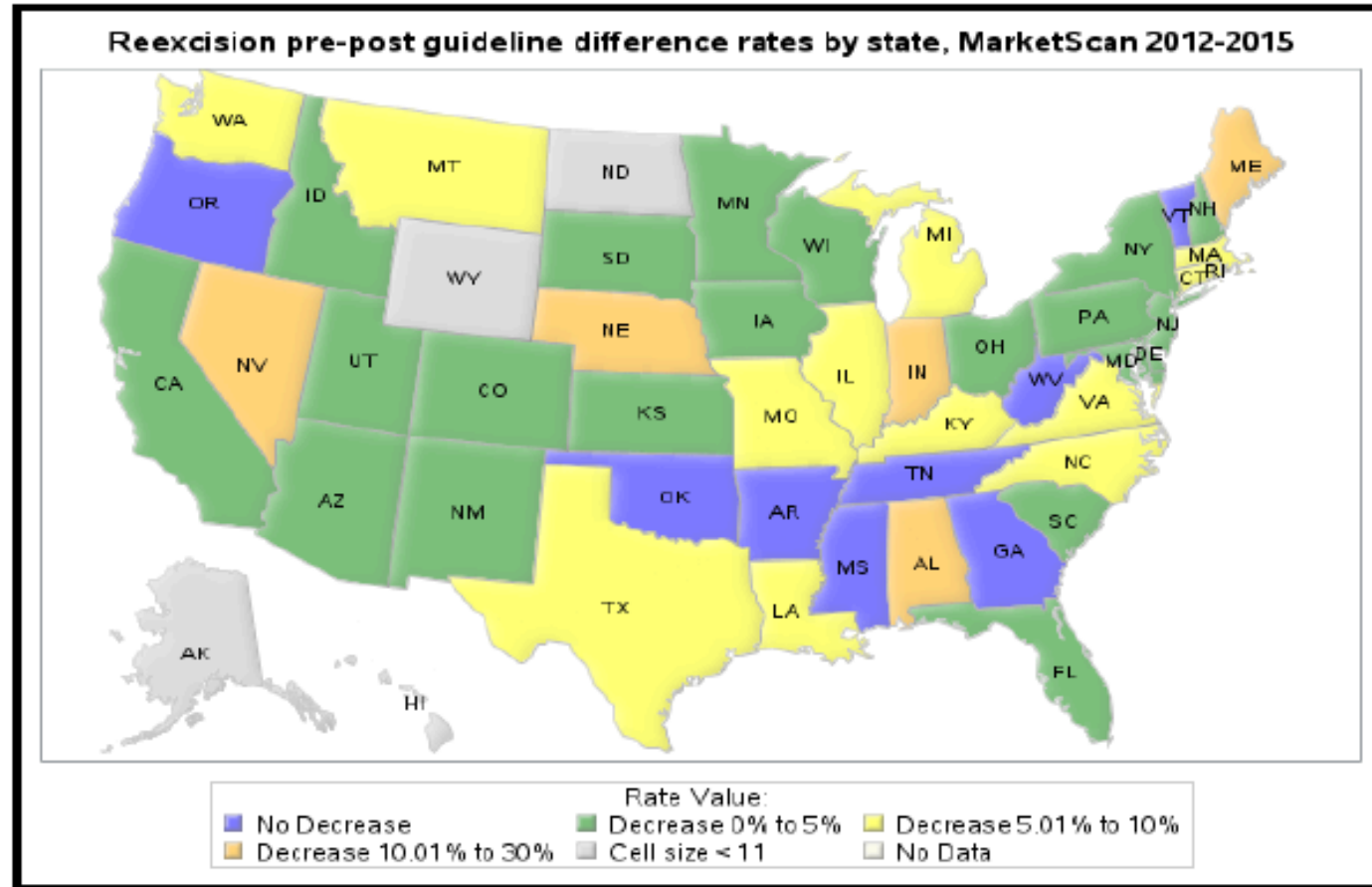
- Patients undergoing BCS in post-guideline period had a statistically significant decrease in risk of re-excision

**FIGURE-1. Quarterly re-excision rates**



# Significant geographical variability by state in re-excision rates

**FIGURE-2. Decrease in the re-excision rate by state**



# Conclusions

- In 2014 there were approximately 113,894 BCS nationwide. A decrease in re-excision rate of 3.7% corresponds to 4,214 fewer unnecessary procedures just in one year.
- The comorbidity and cost associated with this reduction has profound implications for patients and for the health care system

# Appropriate margins for breast cancer conserving surgery in patients with early stage breast cancer- A meta-analysis

Frank Vicini et al

- Wider margins may reduce local recurrence but might increase morbidity and cost
- Previous meta-analysis concluded wider margins unlikely to have substantial benefit
- Current SSO-ASTRO guideline “no tumor on ink”
- Updated meta-analysis
  - Additional studies
  - Evaluate different margin definition models

# Methods

- Systematic literature review 1995-2016
- Inclusion criteria
  - Minimal f/u 50 mo
  - Pathologic margin
  - Local recurrence reported
- 38 studies/55,302 patients
  - Including 31 of 33 previous studies included
  - Additional 20,000 patients
- Median f/u 7.2 years

# Methods

- Margin definition similar to previous analysis
- Positive = invasive or DCIS at surgical margin
- Negative = no tumor within a specified distance from margin
- Close = no tumor on ink but tumor less than a specified distance from the margin
- Series of models developed based on cut-point desired

# Methods

- Previous analysis
  - Model 1- negative margins compared with close/positive
  - Model 2- comparison of negative, close and positive
- Present analysis
  - Model 1- negative margins compared with close/positive

# Model 1

- Consistent with previous model showing that negative margin associated with lower risk of recurrence

**Crude Local Recurrence- Model 1**

Negative vs close/positive

Margin (mm) and status	Number of studies providing data	Total number of observations	Total number of LR	LR %
≤0 >0	28 38	3416 46178	352 1758	<b>10.3</b> <b>3.8</b>
≤1 >1	29 27	3988 33565	405 1161	<b>10.2</b> <b>3.5</b>
≤2 >2	32 23	5978 31757	519 1053	<b>8.7</b> <b>3.3</b>
≤5 >5	38 8	9781 19059	659 604	<b>6.7</b> <b>3.2</b>

Benefit to wider margins observed with greatest difference (benefit) seen at 1 mm (6.7%), but similar rates of recurrence with all negative margin definitions (3.2-3.8%)

Compare:	At or below margin classifications	Above margin classifications
≤0 mm versus >0 mm	≤0	> 0, >1, >2, >3, >5, ≥5; 0-1, 0-2, 0-3, 0-5, 1-2, 2-5
≤1 mm versus >1 mm	≤0, ≤1, 0-1	>1, >2, >3, >5, ≥5; 1-2, 2-5
≤2 mm versus >2 mm	≤0, ≤2, <1; 0-1, 0-2, 1-2	>2, >3, >5, ≥5; 2-5
≤5 mm versus >5 mm	≤0, ≤2, ≤3, ≤5, <1, <5; 0-1, 0-2, 0-3, 0-5, 1-2, 1-5, 2-5	>5

# Model 2

- Looking at range of margins a wider margin further reduced the local recurrence and was validated on MVA
- Margin range
  - >0-2 mm 7.2 % OR: 0.56  $p<0.001$
  - 2-5 mm 3.6% OR: 0.44  $p<0.001$
  - >5 mm 3.2% OR: 0.32  $p<0.001$
- Multivariate Analysis: margin width only significant variable

Range	Margin range classifications
$\leq 0\text{mm}$	$\leq 0$
0-2 mm	0-1, 1-2, 0-2
2-5 mm	2-5
>5 mm	>5

# Model 3 negative, close or positive

- Negative/Close/Positive
  - 1mm: 8.0%/13%/14%
  - 2mm: 3.6%/5.5%/9.5%
  - 5mm: 2.9%/4.1%/12.8%
- Odds Ratio for LR
  - Close vs Neg: 1.58
  - Positive vs Neg: 2.49
  - **2mm vs 1mm: 0.50**
  - **5mm vs 1mm: 0.40** (both of these different from previous)

# Conclusions

- Limitations of meta-analysis
- However: data suggests having a margin width beyond no tumor on ink may further reduce rates of local recurrence
- Further prospective studies required to validate appropriate margin width
- Question?? Should we achieve a 1-2 mm margin as compared to no tumor on ink?

Axillary dissection vs no dissection in patients with cT1-T2 N0 breast cancer and micrometastases only in the sentinel node: 10 yr results IBCSG 23-01 Galimberti et al

- 5 yr results of 23-01 and 10 yr results of Z-0011 showed that for pts with moderate axillary involvement AD provided no advantage for OS or DFS with low axillary failure rate
- These data represent 10 yr update for 23-01 with 83% follow-up completed

# Design

- Prior to amendment in 2006
  - Tumor  $\leq$  3cm, unicentric, one micromet ( $<2\text{mm}$ ), no ECE or macromet
- After amendment
  - Tumor  $\leq$  5cm, uni or multicentric, one or more micromet in sentinel node
- Patients randomized to axillary dissection or no dissection

# Statistics

- Primary endpoint: invasive disease-free survival (DFS)
- Secondary endpoint: overall survival, incidence of re-appearance of tumor in un-dissected axilla
- The non-inferiority margin was defined as a DFS hazard ratio of  $< 1.25$
- 934 patients randomized

# Patient and Tumor Characteristics

Characteristic	AD (n=464)	No AD (n=467)	Total (n=931)
Age, years; median (range)	53 (28–81)	54 (26–81)	54 (26–81)
Menopausal status			
Pre	204 (44%)	207 (44%)	411 (44%)
Post	260 (56%)	260 (56%)	520 (56%)
Pre-op sentinel node biopsy			
No	287 (62%)	286 (61%)	573 (62%)
Yes	177 (38%)	181 (39%)	358 (38%)
Sentinel node disease			
≤ 1 mm	323 (70%)	320 (69%)	643 (69%)
1.1–2.0 mm	131 (28%)	135 (29%)	266 (29%)
>2 mm	10 (2%)	11 (2%)	21 (2%)
Unknown	0	1 (<1%)	1 (<1%)

# Patient and Tumor Characteristics

Characteristic	AD (n=464)	No AD (n=467)	Total (n=931)
<b>Tumor size</b>			
<2 cm	316 (68%)	322 (69%)	638 (69%)
2 cm to 2.9 cm	106 (23%)	112 (24%)	218 (23%)
≥3 cm	35 (8%)	28 (6%)	63 (7%)
Unknown	7 (2%)	5 (1%)	12 (1%)
<b>Tumor grade</b>			
Grade 1	118 (25%)	90 (19%)	208 (22%)
Grade 2	214 (46%)	241 (52%)	455 (49%)
Grade 3	129 (28%)	135 (29%)	264 (28%)
Unknown	3 (<1%)	1 (<1%)	4 (<1%)

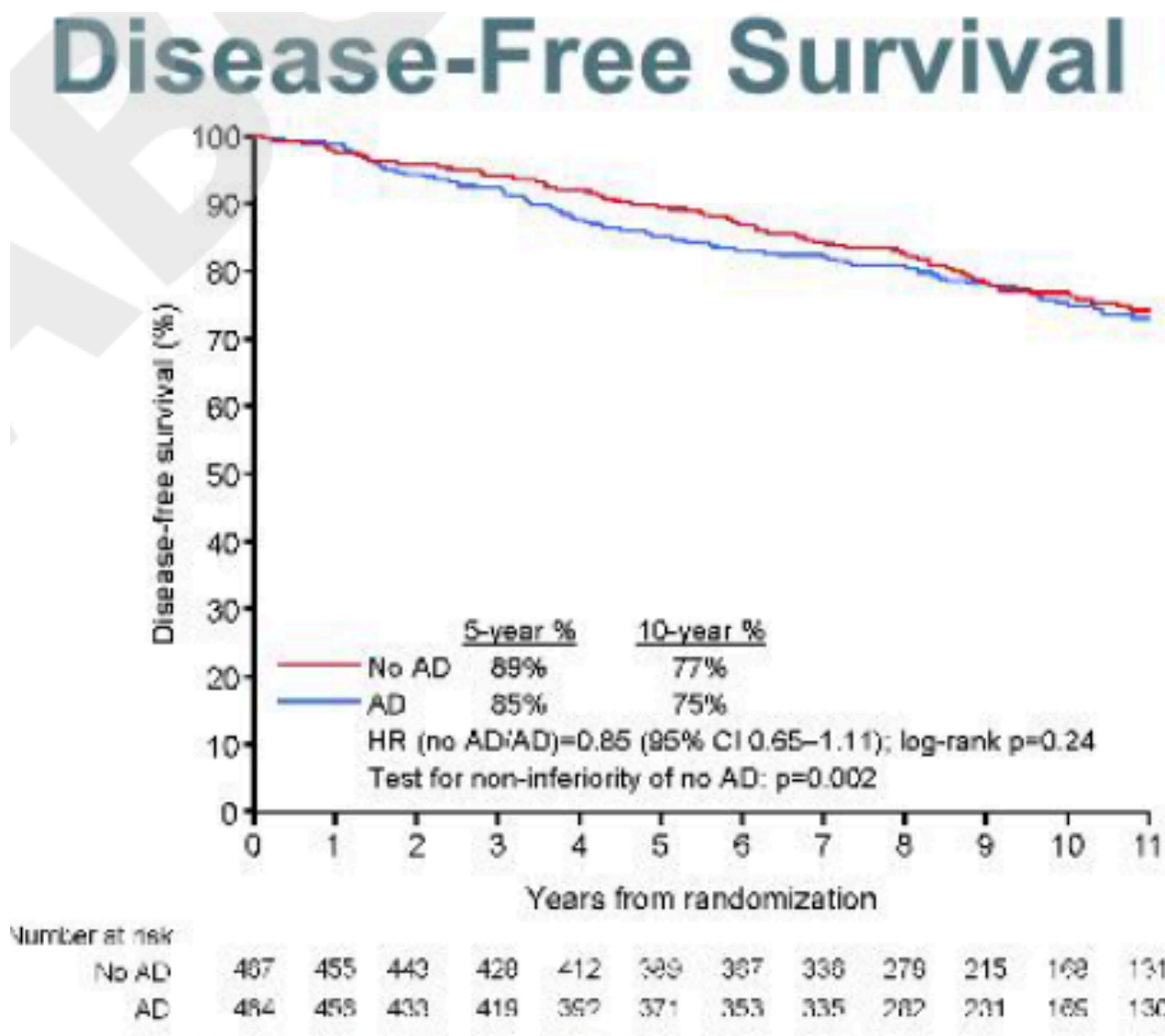
# Patient and Tumor Characteristics

Characteristic	AD (n=464)	No AD (n=467)	Total (n=931)
<b>ER status</b>			
Negative	51 (11%)	40 (9%)	91 (10%)
<b>Positive</b>	<b>409 (88%)</b>	<b>425 (91%)</b>	<b>834 (90%)</b>
Unknown	4 (<1%)	2 (<1%)	6 (<1%)
<b>PgR status</b>			
Negative	108 (23%)	115 (25%)	223 (24%)
<b>Positive</b>	<b>352 (76%)</b>	<b>350 (75%)</b>	<b>702 (75%)</b>
Unknown	4 (<1%)	2 (<1%)	6 (<1%)

Treatment	AD (n=464)	No AD (n=467)	Total (n=931)
Type of breast surgery			
Conservative	420 (91%)	425 (91%)	845 (91%)
Mastectomy	44 (9%)	42 (9%)	86 (9%)
Radiotherapy (BCS)			
No	10/420 (2%)	12/425 (3%)	22 (3%)
Yes	410/420 (98%)	413/425 (97%)	823 (97%)
Radiotherapy in mastectomy			
No	42/44 (95%)	39/42 (93%)	81 (94%)
Yes	2/44 (5%)	3/42 (7%)	5 (6%)

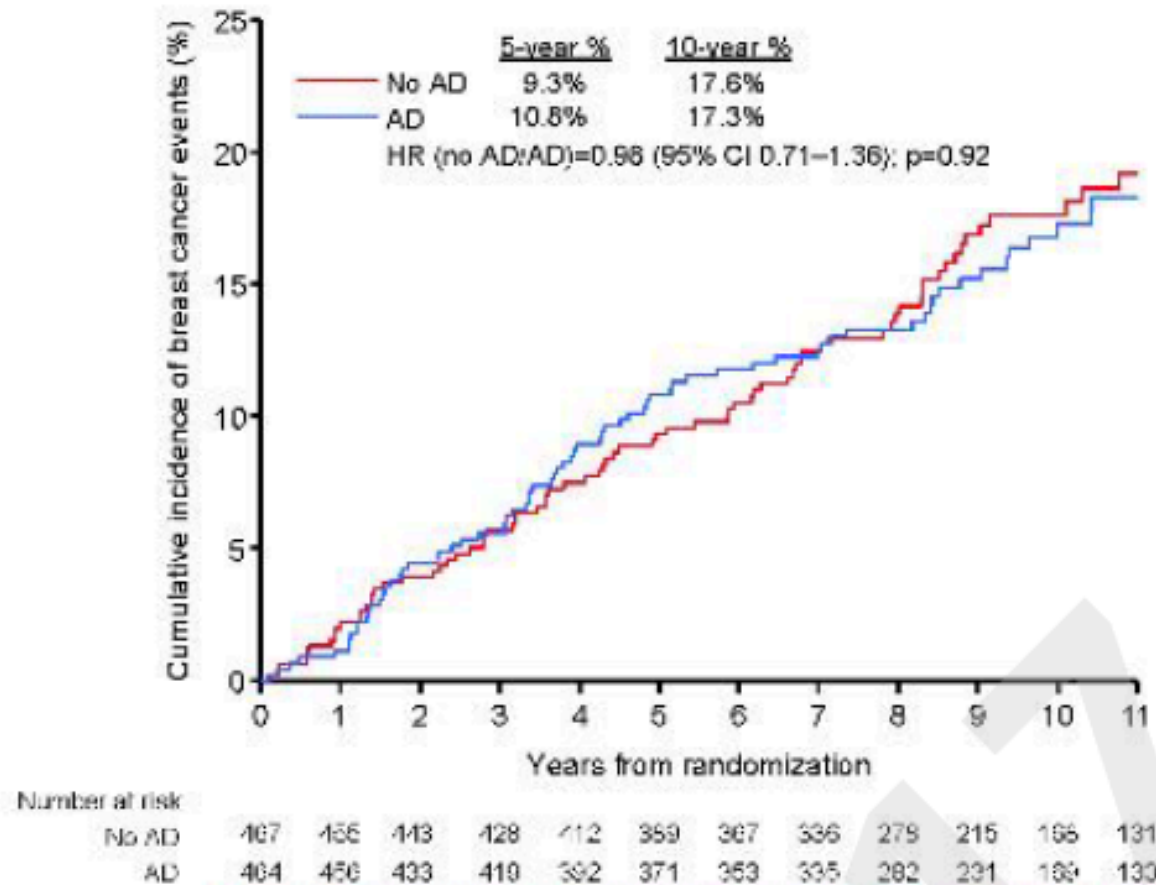
Treatment	AD (n=464)	No AD (n=467)
Any systemic therapy	441 (95%)	451 (97%)
Hormonal therapy only	292 (63%)	315 (67%)
Chemotherapy only	42 (9%)	33 (7%)
Combination therapy	107 (23%)	103 (22%)

# Results



# Results

## Cumulative Incidence of Breast Cancer Events



# Results

## Disease-Free Survival Events and Deaths

	AD (n=464)		No AD (n=467)	
<b>Total DFS events</b>	<b>117</b>	<b>25.2%</b>	<b>101</b>	<b>21.6%</b>
<b>Breast cancer events</b>	<b>75</b>	<b>16.2%</b>	<b>74</b>	<b>15.8%</b>
Local	13	2.8%	14	3.0%
Regional	3	0.6%	9	1.9%
<b>Ipsilateral axillary events</b>	<b>2</b>	<b>0.4%</b>	<b>8</b>	<b>1.7%</b>
Distant	47	10.1%	41	8.8%
Contralateral breast	12	2.6%	10	2.1%
<b>Non-breast cancer events</b>	<b>42</b>	<b>9.1%</b>	<b>27</b>	<b>5.8%</b>
Second (non-breast) primary	23	5.0%	17	3.6%
Death without prior cancer event	2	0.4%	6	1.3%
Death with unknown cancer status	17	3.7%	4	0.9%
<b>Deaths</b>	<b>58</b>	<b>12.5%</b>	<b>45</b>	<b>9.6%</b>

# Results

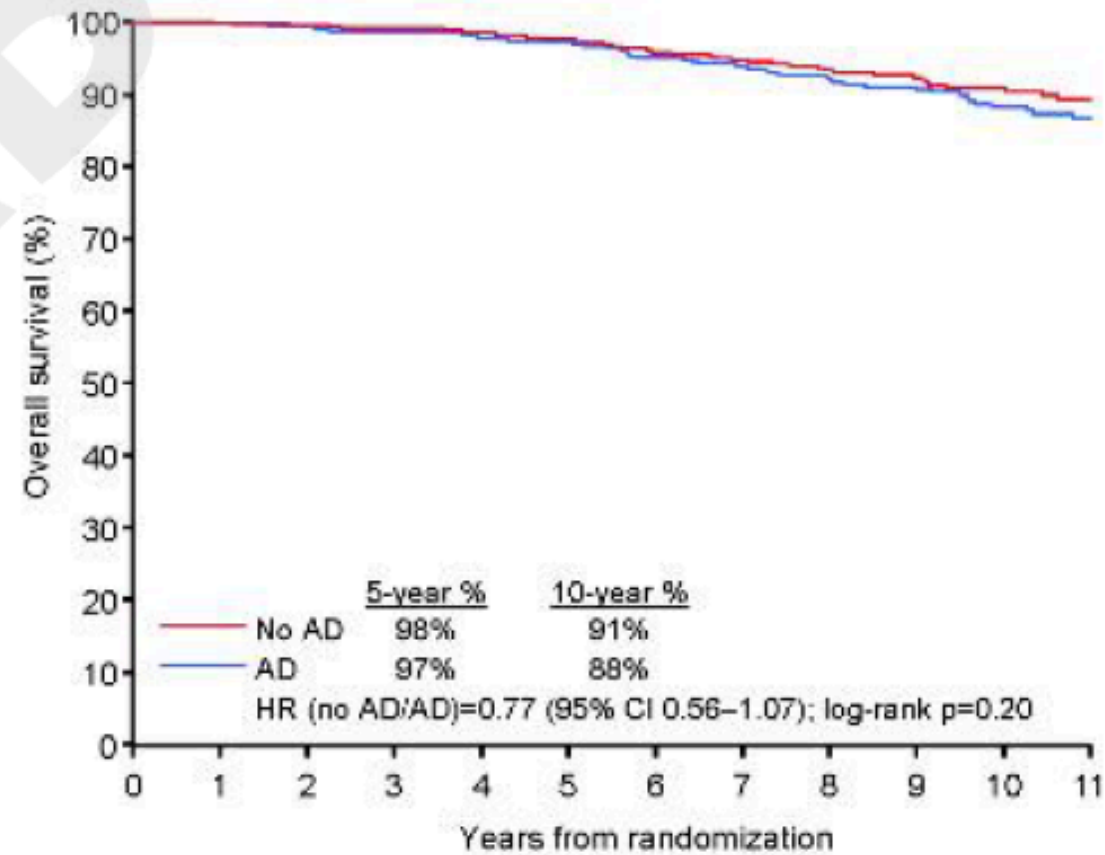
## Disease-Free Survival Events According to Type of Surgery

	Mastectomy (n=86)		Breast conservation (n=845)	
Total DFS events	23 (26.7%)		195 (23.1%)	
Breast cancer events	17 (19.8%)		132 (15.6%)	
Ipsilateral axillary events	2		8	
	AD	No AD	AD	No AD
	1 (1.2%)	1 (1.2%)	1 (0.1%)	7* (0.8%)
Non-breast cancer events	6 (7.0%)		63 (7.5%)	
Deaths	14 (16.3%)		89 (10.5%)	

\*Five received intraoperative radiotherapy

# Results

## Overall Survival



Number at risk

No AD	467	454	460	451	441	424	405	375	314	257	197	158
AD	464	451	456	446	438	422	401	379	319	264	199	156

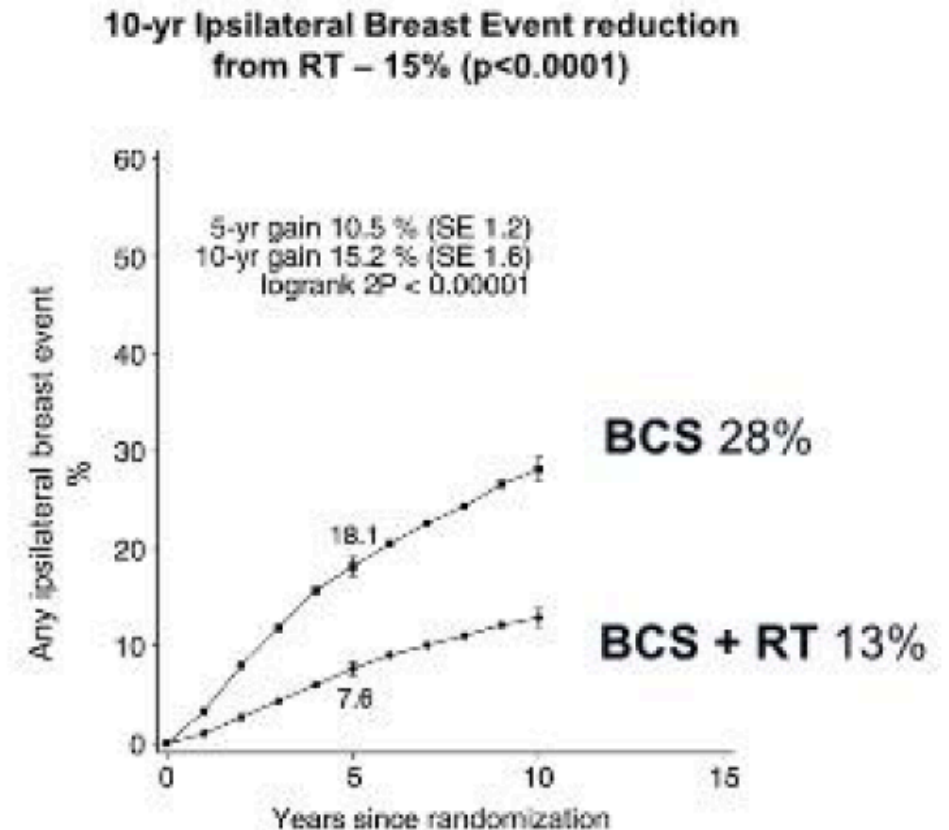
# Summary

- After median follow-up 9.8 yrs
  - No difference between groups for DFS or OS
  - Rate of failure in no AD was low at 1.7%
- Findings are fully consistent with Z011
- We also suggest that non-AD is acceptable treatment in patients scheduled for mastectomy
- No AD is now standard treatment in early breast cancer when the SN is only minimally involved

# A Validation of a DCIS biological risk profile in a randomized study for radiation therapy SweDCIS

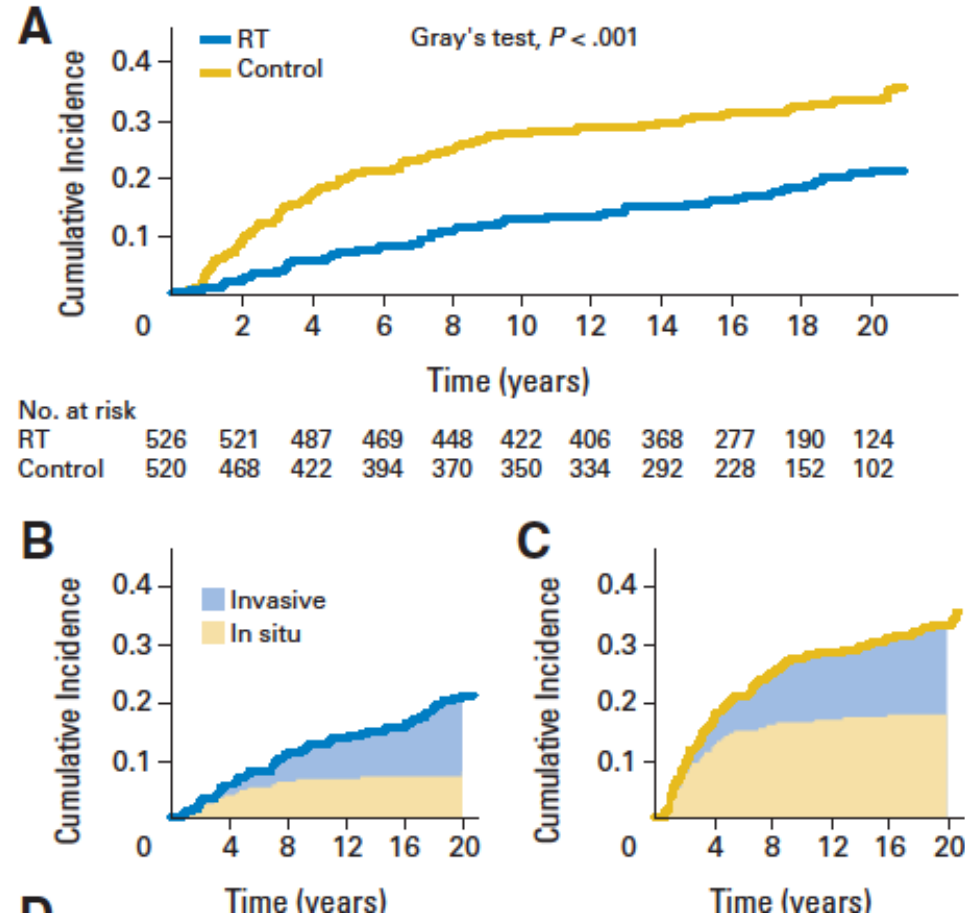
Warnberg, et al

- DCIS randomized trials 2010 EBCTCG Overview
- Approximate 50% reduction in risk of recurrence with XRT
- 50% reduction independent of size, age or grade



# SweDCIS Randomized Trial

- Eligibility criteria
  - Primary, pure DCIS
  - 1987-1999, BCS +/- XRT
- Recurrences
  - 129 in situ
  - 129 invasive

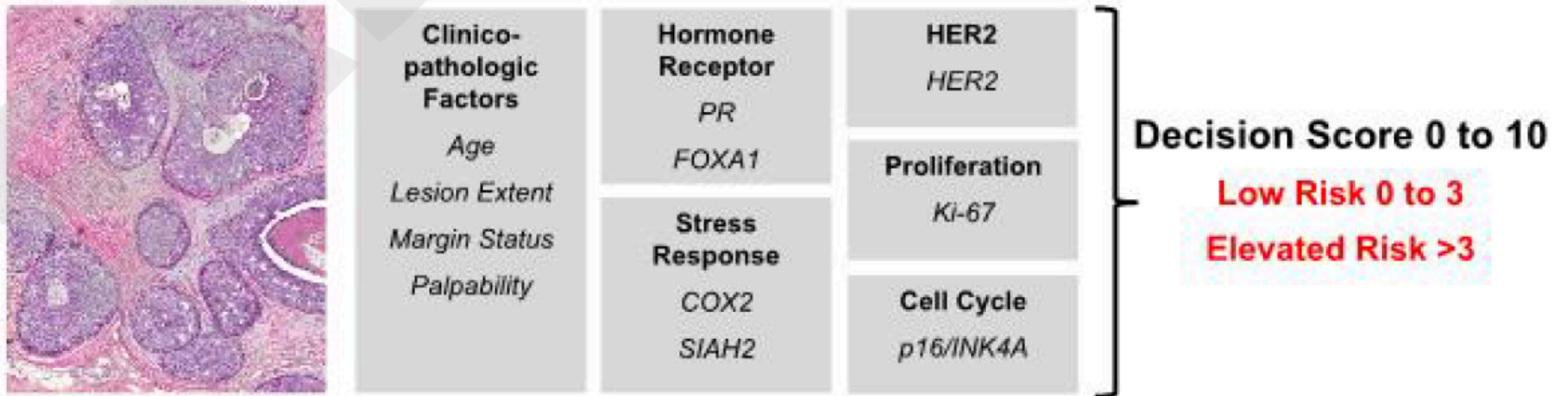


# Validation of Biological Risk Signature

- Assess correlation between
  - Total ipsilateral events/invasive
  - Decision Score
    - Low risk group and RT
    - Elevated risk group and RT
- Multivariate Interaction Analysis

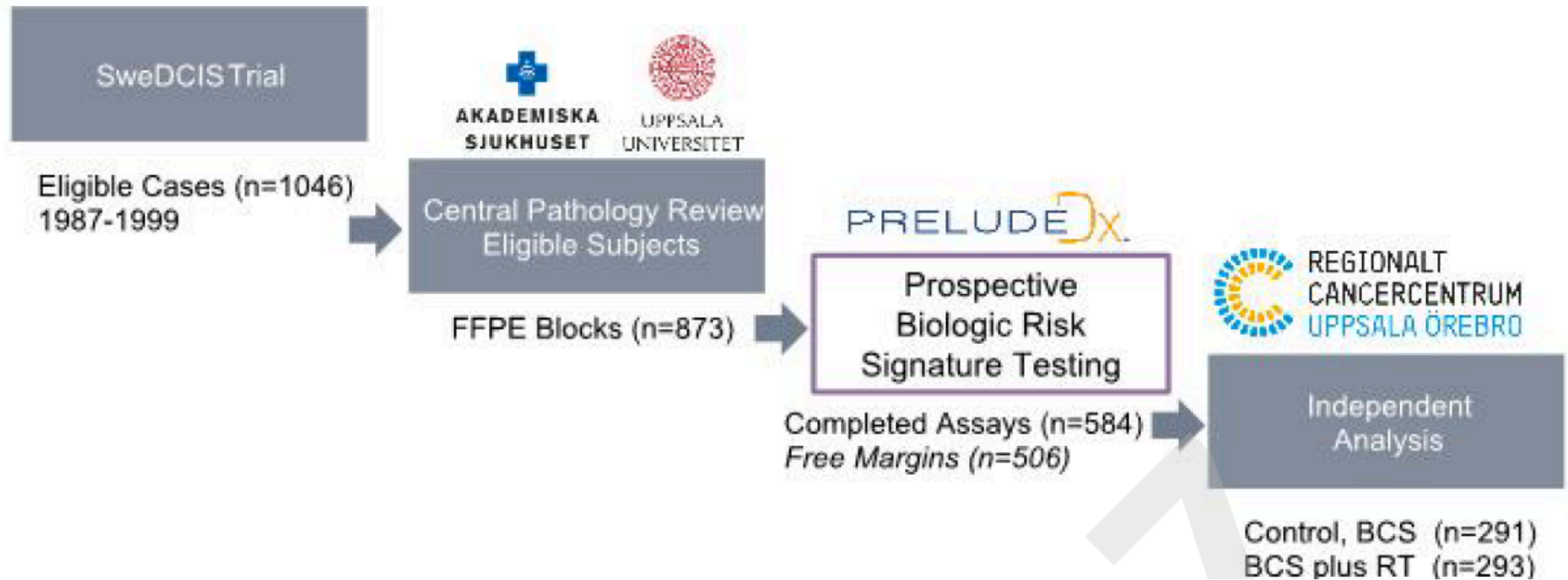
# Who will benefit from RT?

## Biologic Risk Signature



- Previously completed validations
  - Multiple cross validation, Uppsala Univ and UMass (n=526) 2015
  - Independent validation, Kaiser Permanente NW (n=455) 2016

# Validation SweDCIS



# SweDCIS Clinical Risk Factors

	<b>Complete Assay</b>	<b>Overall Trial</b>
Characteristic	Count (%)	Count (%)
Age <50 years	140/584 (24%)	252/1046 (24%)
Size ≥10 mm	424/584 (73%)	709/935 (76%)
Margins: Involved	78/584 (13%)	112/952 (12%)
Treatment Year <1995	331/584 (57%)	602/1046 (58%)
BCS plus RT	293/584 (50%)	526/1046 (50%)

# SweDCIS: RT Benefit at 10 yrs

Complete Assay Data w/ Clear Margins, n=506

No significant RT benefit in Low Risk group

## Multivariate Interaction Analyses

Characteristic	n	Invasive Breast Cancer (IBC)		Total IBE (DCIS+IBC)	
		HR	p-value	HR	p-value
Low Risk (DS≤3)	243	0.84 [0.32 to 2.22]	0.70	0.48 [0.24 to 0.97]	0.04
Elevated Risk (DS>3)	263	0.24 [0.08 to 0.76]	0.012	0.31 [0.17 to 0.59]	<0.001

RT benefit larger than expected 50%

# SweDCIS compared to Cross-validation

clear margins

- RT benefit at 10 yrs, multivariate interaction analyses

Low Risk (DS≤3)		Invasive Breast Cancer (IBC)		Total IBE (DCIS+IBC)	
Study	n	HR	p-value	HR	p-value
SweDCIS	243	0.84 [0.32 to 2.22]	0.70	0.48 [0.24 to 0.97]	0.04
UUH/UMASS (Cross-Validation)	196	0.76 [0.23 to 2.53]	0.65	0.50 [0.21 to 1.18]	0.12

Elevated Risk (DS>3)		Invasive Breast Cancer (IBC)		Total IBE (DCIS+IBC)	
Study	N	HR	p-value	HR	p-value
SweDCIS	263	0.24 [0.08 to 0.76]	0.012	0.31 [0.17 to 0.59]	<0.001
UUH/UMASS (Cross-Validation)	278	0.32 [0.13 to 0.77]	0.011	0.26 [0.14 to 0.47]	<0.001

# Conclusion

- Biological risk signature (Decision Score) correlated to risk (**prognostic**)
- Biological risk signature **predicted** RT benefit
  - Not all patient groups have equivalent RT benefit
  - Low risk patients no RT benefit to prevent Invasive recurrence
  - Elevated risk patients substantial benefit nearly 2x expected

# Local recurrence and salvage lumpectomy

Sellam et al

## Local Recurrence of Breast Cancer: Salvage Lumpectomy as a Safe Option for Local Treatment

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### Abstract

The best local management for breast cancer recurrence following conservative treatment for breast cancer (breast conserving surgery followed by radiation therapy) continues to be an open question. In this study, we compared patient outcomes after salvage lumpectomy vs. patients who underwent mastectomy for ipsilateral breast tumor recurrence (IBTR).

### Objectives

Between 1987 and 2014 we identified 121 patients with pT0-2, N0-3, M0 breast cancer who had breast conserving surgery and radiation as their primary treatment, and subsequently had IBTR (unifocal only, eligible for re-lumpectomy).

At first recurrence, 47 patients (39%) underwent salvage lumpectomy (SL) and 74 (61%) salvage mastectomy (SM).

Median F/U was 14 years (2-30) from first breast cancer diagnosis.

### Results

The women who opted for SM as salvage treatment for their recurrence previously had more ALND (58% vs. 33%,  $p=0.023$ ) and received neoadjuvant chemotherapy more frequently (20% vs. 4%,  $p=0.022$ ) at their first breast cancer diagnosis. The median DFI at 1<sup>st</sup> recurrence for SL and SM group was 12 and 7 years, respectively ( $p=0.011$ ).

	Salvage Lumpectomy Patients, n (%)	Salvage Mastectomy Patients, n (%)	P-value
Mean Age at 1 <sup>st</sup> diagnosis	52.2 y	46.9 y	0.006
Tumor size at 1 <sup>st</sup> surgery	1.26 cm	1.67 cm	0.722
Tumor size at 2 <sup>nd</sup> surgery	1.13 cm	1.52 cm	0.013
RT post 2 <sup>nd</sup> surgery	16 (34%)	20 (27%)	0.154
Pathologic G 1 <sup>st</sup> surgery			0.046
Grade 1	14 (29.7%)	12 (16.9%)	
Grade 2	21 (44.6%)	30 (42.2%)	
Grade 3	6 (12.7%)	22 (30.9%)	
DCIS related	6 (12.7%)	7 (9.8%)	
IDC pathology 1 <sup>st</sup> surgery	(74%) 35	(83%) 59	0.761
ER 1 <sup>st</sup> surgery - Positive	(74%) 28	(60%) 40	0.202
HER2 1 <sup>st</sup> surgery Positive	(24%) 9	(26%) 17	0.819
True Recurrence	(46.8%) 22	(39.4%) 28	0.395

- ❖ For the SL and SM cohorts, 8 and 10 patients (17%, 13.5%,  $p=0.22$ ) respectively, developed subsequent local recurrence as a 3<sup>rd</sup> event. The median DFI between 2<sup>nd</sup> and 3<sup>rd</sup> recurrence for SL and SM was 6.5 and 15.5 years, respectively ( $p=0.081$ ).
- ❖ In a univariate analysis, age at 1<sup>st</sup> diagnosis, T stage at 1<sup>st</sup> diagnosis and 1<sup>st</sup> recurrence, number of dissected/+ve LN, grade, type of axillary surgery, neo/adjuvant chemo and HER2 status (both in primary and at recurrence), had no effect on 2<sup>nd</sup> recurrence occurrence for both groups.
- ❖ In MVA, undergoing SL had higher chances of having a 2<sup>nd</sup> recurrence (3<sup>rd</sup> event),  $p=0.020$ . Having re-irradiation following SL did not protect against 2<sup>nd</sup> recurrence (3<sup>rd</sup> event,  $p=0.42$ ).
- ❖ At a median follow-up of 14 years, 95.7% of SL patients are alive, NED, 85% are mastectomy free. 84% of patients who opted for SM are alive, NED.32

### Conclusion

Salvage lumpectomy following IBTR, while associated in MVA with higher second LR rate than SM, is not associated with inferior survival. With survival >95% at 14 years in the SL cohort, salvage lumpectomy with or without re-radiation, in a selected population (unifocal T), represents an acceptable treatment option for patients in order to delay time to mastectomy and keep the original breast without reducing BC survival. Both options should be discussed prior to any surgical decision.

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Code

# Salvage lumpectomy after recurrence

- Between 1987 and 2014 121 patients identified
- pT0-2, N0-3
- BCS and XRT
- In breast true recurrence (unifocal) eligible for re-lumpectomy
- 47 (39%) underwent salvage lumpectomy
- 74 (61%) underwent salvage mastectomy
- Median f/u 14yrs from first diagnosis

	Salvage Lumpectomy	Salvage Mastectomy	
	Patients, n (%)	Patients, n (%)	P-value
Mean Age at 1 <sup>st</sup> diagnosis	52.2 y	46.9 y	0.006
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# Results

- For SL and SM cohorts, 8 and 10 pts (17%, 13.5%  $p=0.2$ ) respectively, developed subsequent local recurrence as 3<sup>rd</sup> event
- Median DFI between 2<sup>nd</sup> and 3<sup>rd</sup> event 6.6 (SL) and 15.5yrs (SM)
- Median follow-up 14yrs, 95.7% of SL patients are alive NED and 85% are mastectomy free

# Conclusions

- In a selected population salvage lumpectomy is an acceptable treatment for women who have recurrence following BCS and XRT

Thank you