

Surgical Therapy: Sentinel Node Biopsy and Breast Conservation

Stephen B. Edge, MD Professor of Surgery and Oncology Roswell Park Cancer Institute University at Buffalo



National Comprehensive

Dr. Roswell Park:

NCCN Cancer Tradition in Cancer Research

- 1898: Founded NY State Laboratory for the Study of Malignant Disease
 - Re-named Roswell Park
 Memorial Institute in 1942
 - RPMI 1640 culture media
- Chair, Dept of Surgery University at Buffalo
- Performing surgery in Niagara Falls when President McKinley shot at the 1901 Pan American Exposition in Buffalo





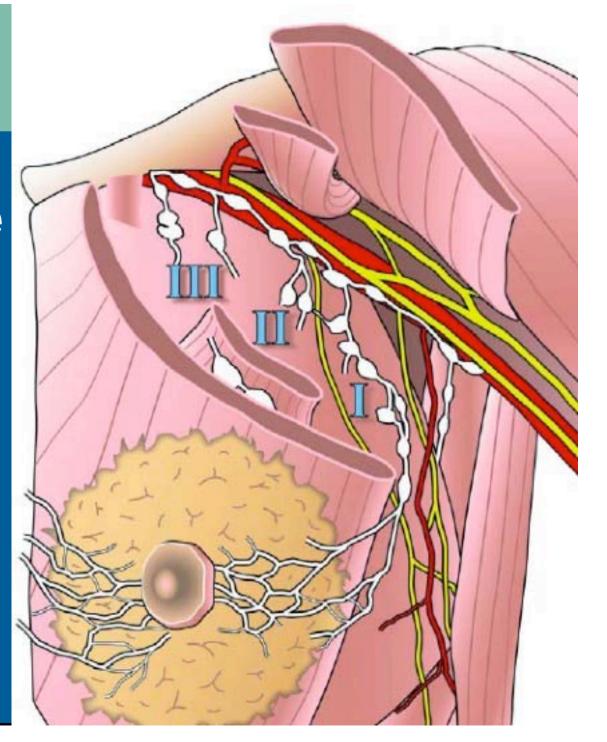
Overview

- Sentinel Node Biopsy
 - Indications for SNB
 - Need for completion axillary dissection
- Breast conservation
 - Techniques for resection of margins
 - Extent of resection with PCT
- Dr. Collins Pathologic evaluation of margins
- Japanese Comment



Sentinel Node Biopsy

Indications





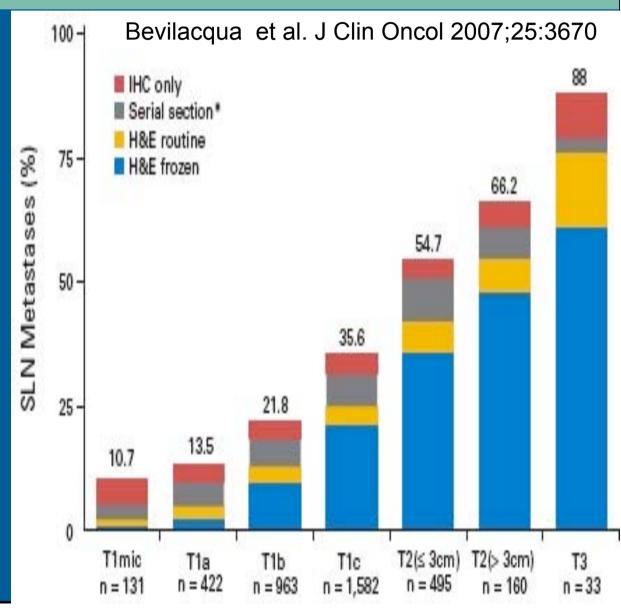
SNB with Clinically Negative Nodes

- Invasive breast cancer
 - Any situation requiring lymph node staging
 - Primary (neoadjuvant chemotherapy)
 - Local recurrence repeat SNB?
- Ductal carcinoma in situ
 - Mastectomy
 - Other indications?



SNB Especially Important with Small Cancers

- Tumors < 1 cm</p>
 - 15% positive nodes by H&E
 - Major impacton use ofchemotherapy





SLNB Indication: Repeat SNB with Recurrence

- Uncertain role of systemic therapy
- Uncertain need for lymph node staging with local recurrence

- Repeat SNB technically possible in women with prior SNB
 - Breast conserving surgery
 - May be possible with prior mastectomy



Repeat Sentinel Node Biopsy

Series	Number	Successful mapping	Drainage outside axilla	Repeat SLN Positive
Moffitt	56	45	1	9 (20%)
John Wayne	6	5	2	0
MSKCC	54	40	3	5 (12.5%)
European Institute	65	63	7	7 (11%)
TOTAL	181	152 (84%)	11 (6%)	20 (13%)

Adapted from Cox et al. JACS 2008;207:57



Lymph Node Surgery with

DCIS



Most "Positive Nodes" with DCIS Only by IHC

Series	N Cases	Positive by H&E	Positive by IHC
Broekhuizen	66	1%	11%
Wilkie	559* All DCIS	1%	5%
Katz	110 High risk	4%	8%
Veronesi	508 All DCIS	2%	1%



Cancer Network

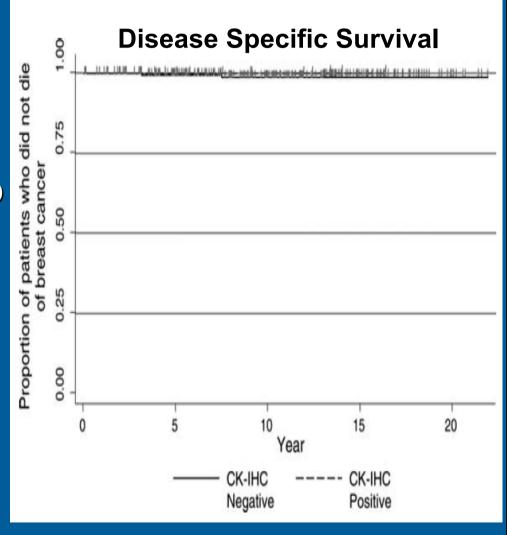
National No survival impact of IHC Comprehensive positive nodes in DCIS

301 pts with DCIS and negative nodes

Median 10 yr follow-up

Cytokeratin IHC on archived blocks

18 / 301 positive by IHC



El-Tamer et al. Ann Surg Onc Disease 2005;12:254

NCCN Guidelines: DCIS

Lumpectomy:

No lymph node surgery

Mastectomy:

Sentinel node biopsy



Lymph Node Surgery with Primary (Neoadjuvant) Chemotherapy



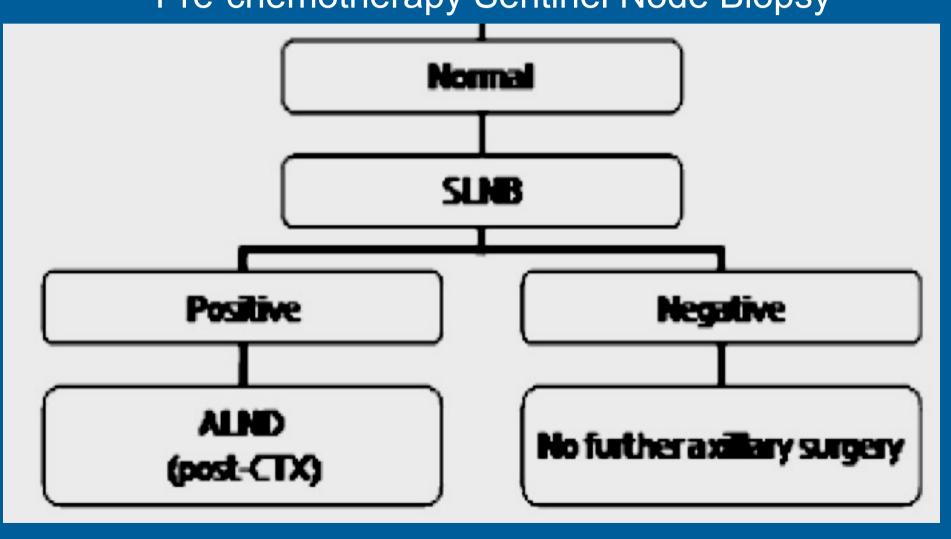
Pre-chemotherapy Sentinel Node Biopsy

	# pts	Negative SNB Before Chemo	Positive SNB Additional Positive Nodes After Chemo
Sabel – Michigan	25	12 (48%)	8 / 13 (60%)
Schrenk - Linz, Austria	21	12 (55%)	6 / 9 (66%)
Cox - Moffitt (LABC)	47	7 (15%)	27 / 40 (67%)
Van Rijk – Netherlands	25	14 (56%)	5 / 11 (45%)

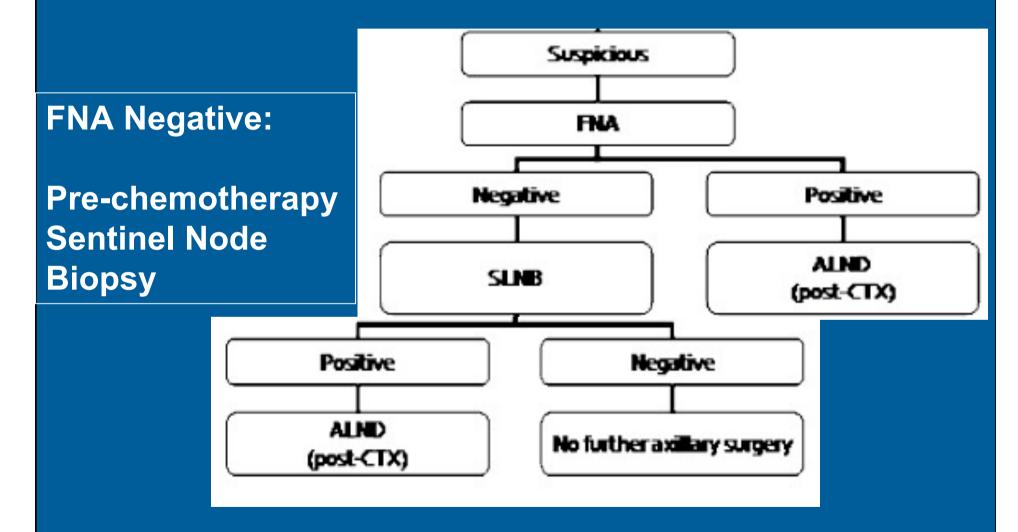


Axillary Management: Clinically / Ultrasound Negative

Pre-chemotherapy Sentinel Node Biopsy







Kilbride KA Annals of Surgical Oncology 2008



Is Axillary Dissection Needed with Positive Sentinel Node?

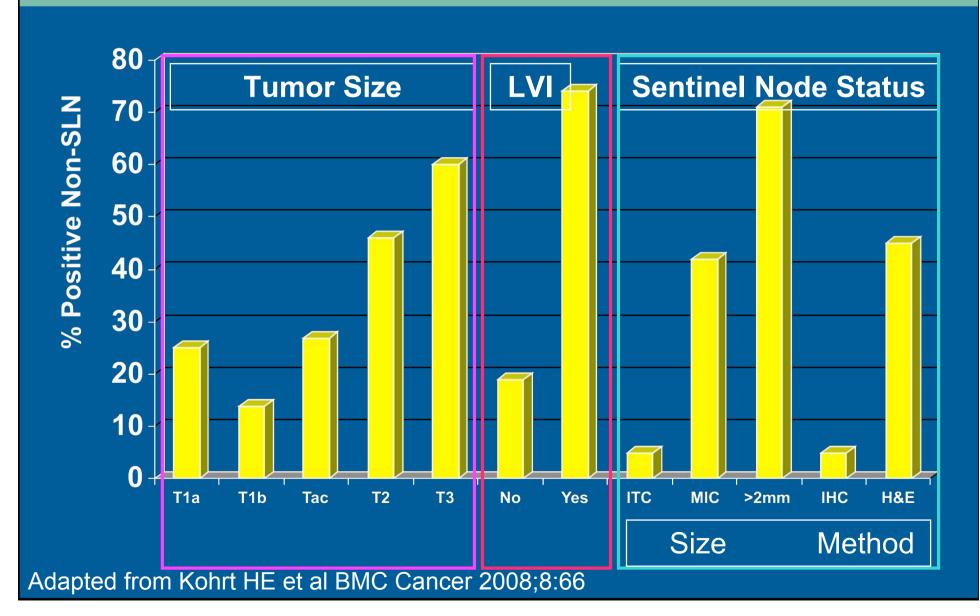


Axillary Dissection with Positive Sentinel Node?

- What is the probability of additional positive nodes?
 - Is there a rate so low that dissection not warranted?
- Therapeutic impact of dissection
 - Do additional positive nodes alter choice of chemotherapy?
 - Control of cancer in axilla: Surgery vs. Radiation

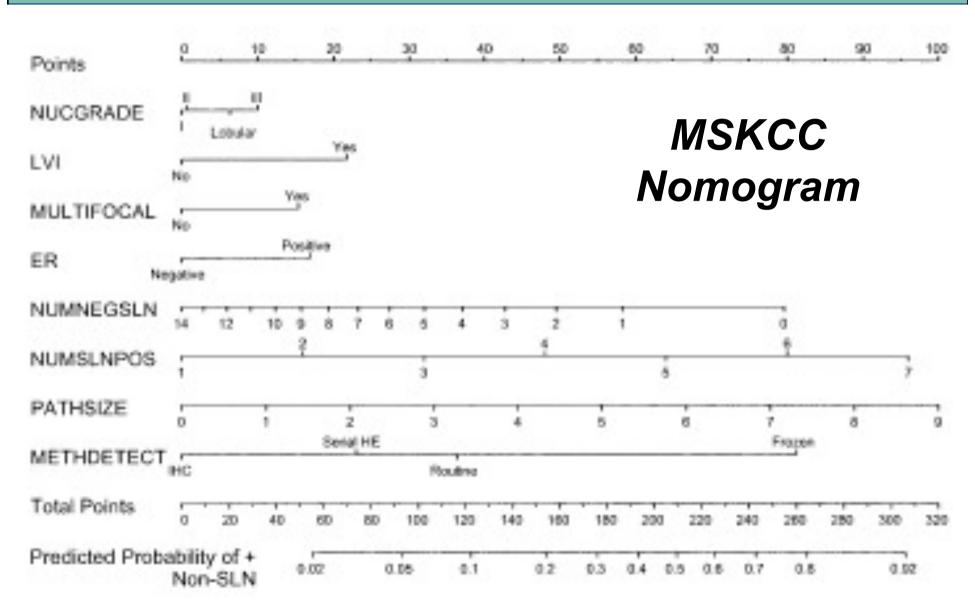


Probability of Additional Positive Nodes





Risk of Additional Involved Nodes with Positive SNB





Omit Axillary Dissection with Positive SNB?

- What risk of additional positive nodes is low enough?
- Most American oncologists perform axillary dissection for any positive nodes

 Major question is in cases of ITC and micrometstases detected by cytokeratin immunohistochemistry



Surgical Margins

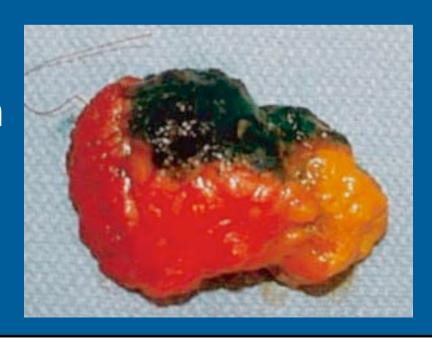


Surgical Margins

1. Techniques

2. Resection after Primary Chemotherapy

3. Pathology Evaluation Dr. Collins





Surgeons' Definition of Negative Margins

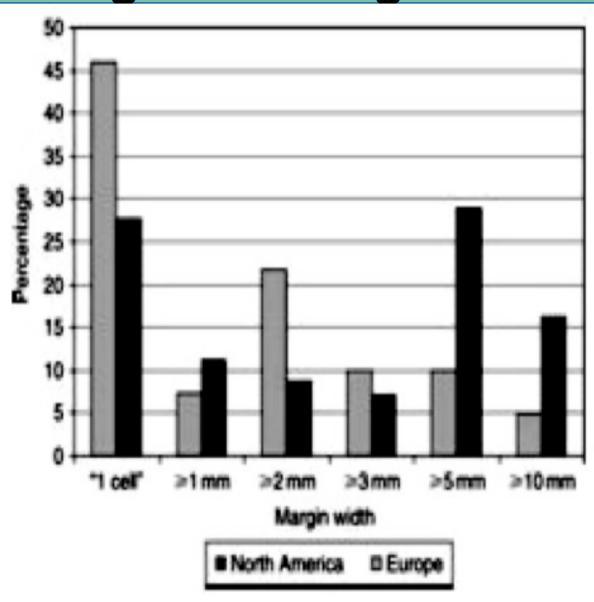
Survey of surgeons

in North America

and Europe on

what is accepted as

"negative margin".





Margin Management

- Careful surgical planning
 - Pre-operative diagnosis of cancer
- Orient specimen
- Specimen mammography
 - Key for calcifications



Techniques for Margin Excision

Primary excision

- Single specimen versus
- Shave margins after primary excision

Re-excision

- Whole cavity versus
- Directed excision of specific margin

Intraoperative evaluation

- Specimen mammography
- Gross
- Microscopic generally NOT performed



Separate Cavity Margin Sampling

- Excision of cancer
- Resection of additional tissue at each of 6 margins

Mass in breast Excision/ lumpectomy (also anterior/ posterior) superior superior total of 6 posterior additional inferior margins anterior inferior

Cao et al. Am J Surg Path 2005;29:1625



Cavity Margin Sampling

- Residual cancer in cavity margin sample in many cases
- Factors associated with residual cancer
 - Extensive intraductal component
 - High grade
 - Extent of margin involvement
- Reduced re-excision by 60%

Primary Lumpectomy Margin	Cavity Margin Sample Contained Residual Cancer
Positive (n=233)	30%
Negative (n=281)	10%

Cao et al. Am J Surg Path 2005;29:1625



National Comprehensive Pre-Surgical Diagnosis Cancer Network Improves Margin Management

- Re-excision after lumpectomy common
- Negative margin more likely with pre-surgical diagnosis - FNA or core biopsy

NCCN study of re-excision

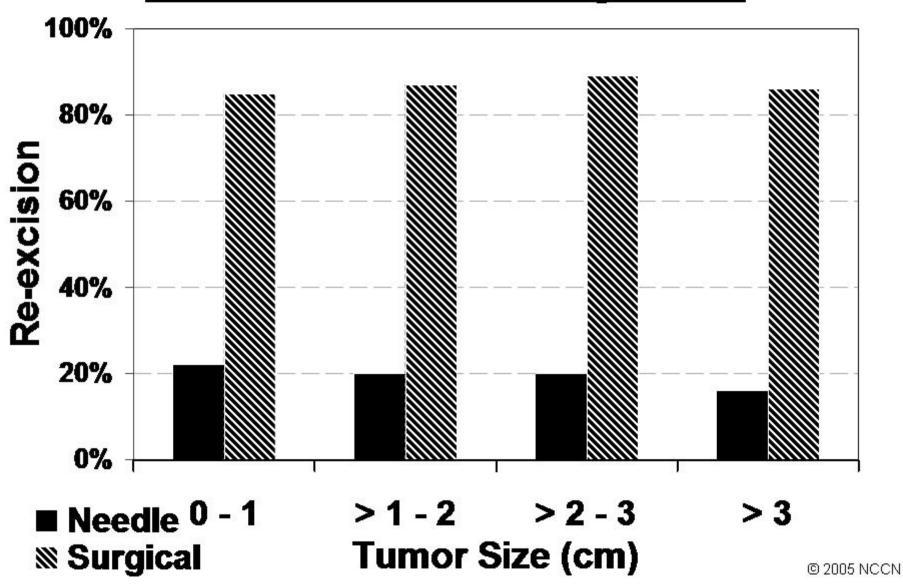
Frequency of re-excision among 6,131 women from 1997 - 2001 based on the type of initial biopsy

Re-excision by Type of Initial Biopsy

Type of Initial Biopsy	N	Percent Re-excision	
Needle	3,481	23%	
Surgical	2,650	92%*	

^{*}Using univariate logistic regression, association between the type of initial biopsy and re-excision is statistically significant (p-value < 0.0001).

Re-excision by Tumor Size for Patients Receiving BCS





Factors Associated with Re-excision

Odds Ratio

Use of surgical biopsy 3.35

Smaller breast 2.7

Lobular histology 1.93

Adjuvant vs. neoadjuvant 2.49

Waljee JF et al. Ann Surg Oncol 2008;15:1297



Re-Excision: Impact on Recurrence

Re-Excision Among 2,770 patients at Fox Chase Overall Re-Excision Rate 60%

Number of	Number of	Local Recurrence		
Re-excisions	patients	5 yr (%)	10 yr (%)	
0	1119	2.5%	5.6%	
1	1514	1.9%	5.7%	
0 and 1	2633	2.1%	5.6%	
2 or more	137	5.5%	10%	

O'Sullivan MA et al. Ann Surg Oncol 2007;14:3133

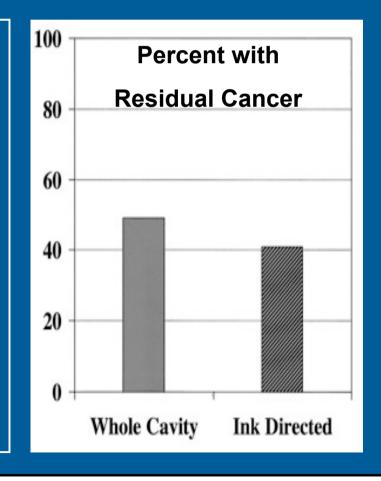


Technique of Re-excision: Prefer Ink-Directed

Whole Cavity versus Ink Directed Resection of Positive Margin

546 lumpectomy
245 (45%) - No re-excision
181 Whole Cavity
120 Directed Resection

- Less tissue removed
- Better cosmetic result
- No difference recurrence





Neoadjuvant Chemotherapy

Defining Extent of Resection

- Use imaging to define extent of cancer prior to and after chemotherapy
- Place marker to allow radiological localization
- Extend surgery around area of original cancer
- Experience and judgment

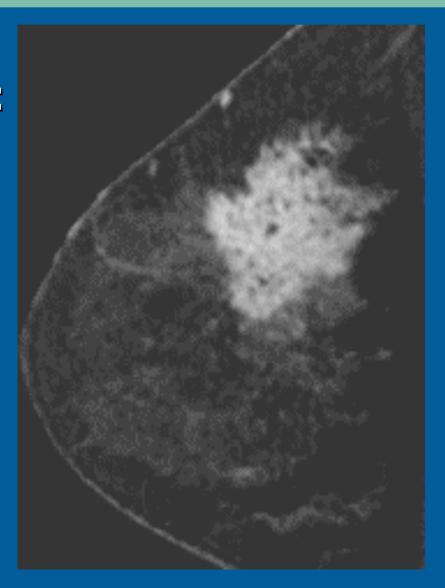


MRI Useful to Define Extent of Cancer

Define size of cancer: supplements mammography

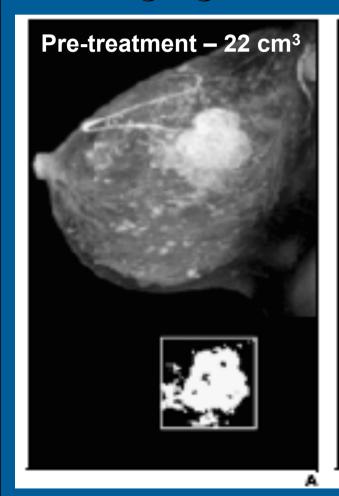
May help define extent of DCIS

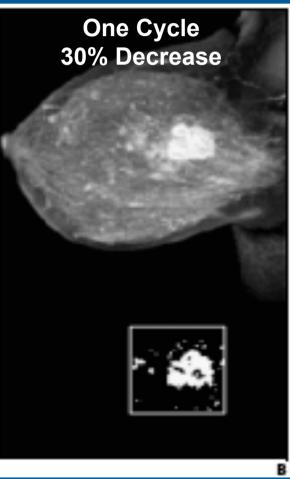
Identify 2nd cancers

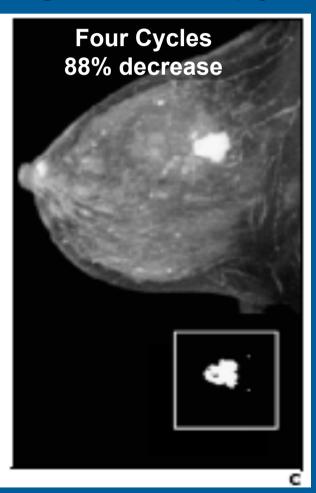




Staging and Response to Pre-surgical Therapy









MRI underestimates residual disease

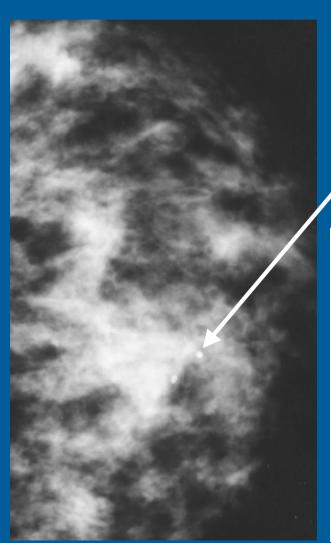
Pathologic	Response by MRI			
Response	CR	PR	NR	Prog
CR	12	0	0	0
PR	10	37	0	0
NR	1	1	7	1
Prog	0	0	0	0

Warren Br J Cancer 2004;90:1349



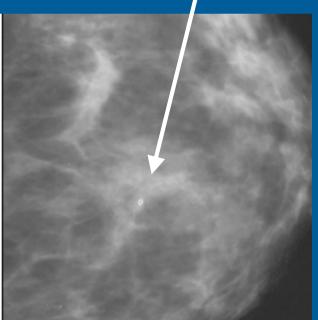
REMEMBER TO PLACE CLIP!!!

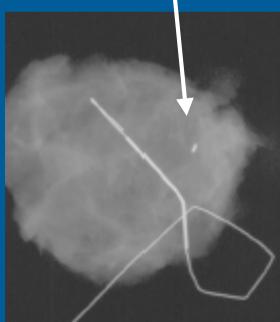
Before Chemotherapy



After Chemo

Specimen |







National Comprehensive Rate of Positive Margin with Neoadjuvant Therapy

Positive Margins:

21% Neoadjuvant

Not neoadjuvant 18%

Factors affecting positive margins:

Lobular cancer 43%

16% **Ductal cancer**

Soucy G et al. J Am Coll Surg 2008;206:1116



Neoadjuvant Therapy: Impact on Extent of Surgery

- Neoadjuvant therapy reduces the extent of surgery
- Does not increase positive margins or reexcision

Neoadjuvant Primary Surgery
Volume 113 cm³ 213 cm³

Re-excision 13% 16%

Boughey JC et al. Ann Surg 2006;244:464

Japanese Experience

Sequential anthracycline / taxane

10% complete pathologic response

38% had lumpectomy

25% with positive margins

Fukutomi T. Breast Cancer 2006;13:147