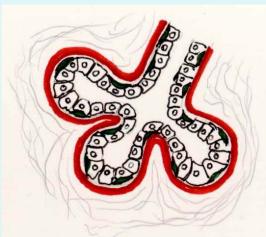
Breast Pathology

Helge Stalsberg MD University Hospital of North Norway



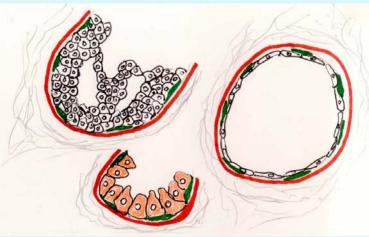
Major classes of breast disease



Normal breast



Carcinoma in situ

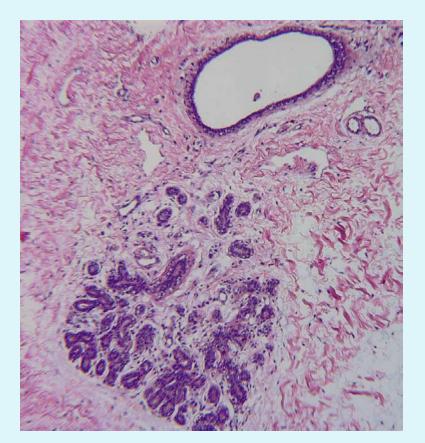


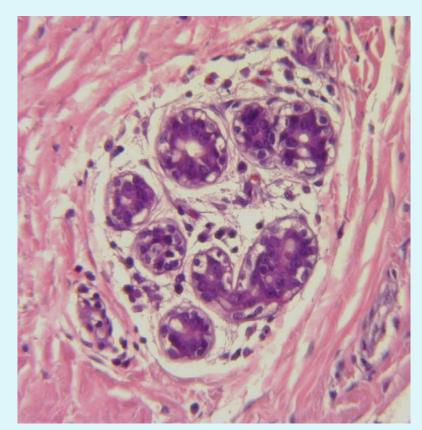
Benign breast disease



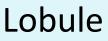
Invasive carcinoma

Normal breast



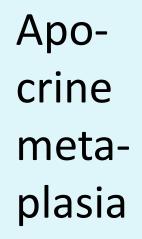


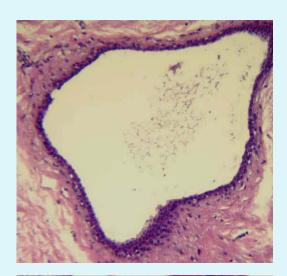
Lobule and duct



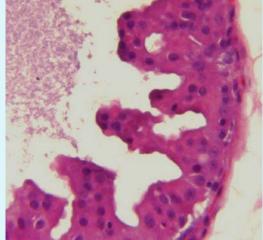
Benign Breast disease: Elements of fibrocystic disease



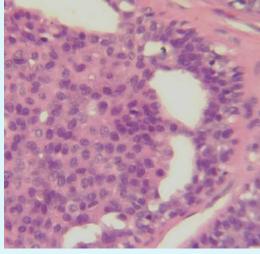


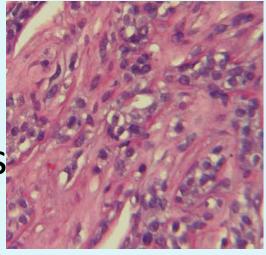


Ductal hyperplasia



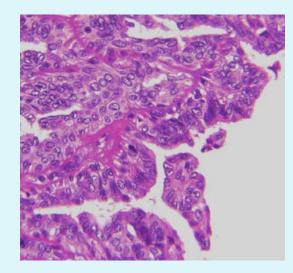
Sclerosing adenosis



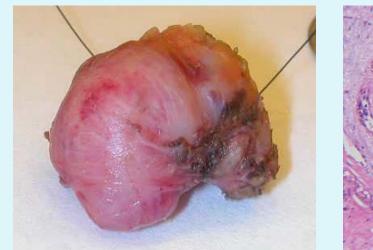


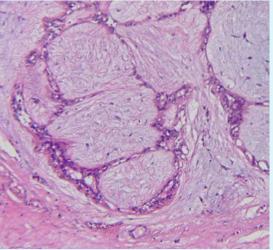
Benign breast disease:

Intraductal papilloma

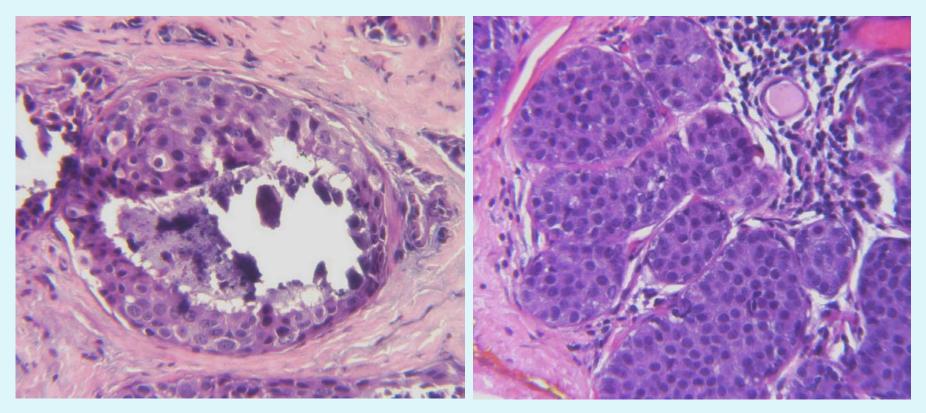


Fibroadenoma





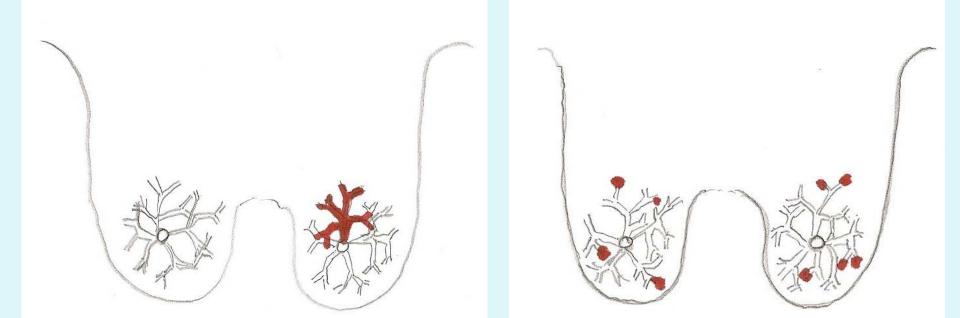
Carcinoma in situ



DCIS

Ductal carcinoma in situ. Lobular carcinoma in situ. LCIS

The distribution of carcinoma in situ



Ductal carcinoma in situ

Lobular carcinoma in situ Invasive carcinoma: The pathology report

- The definite diagnosis
- The local/regional extent of the disease
- Data relevant to prognosis
- Data relevant for the choice of treatment

The Pathology report, Ca. resection

MAIN DIAGNOSIS:

• Local resection (lumpectomy) of breast with invasive ductal carcinoma

OTHER FEATURES:

- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast
- TUMOR GRADE: 2
- <u>RESECTION MARGINS, MINIMUM DISTANCE TO TUMOR</u>: 4 mm from dorsal margin, 32 mm from ventral margin
- <u>TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE</u>: Absent.
- VASCULAR INVASION: Present
- DCIS: Present

DCIS GRADE: 3 DCIS MIN. DIST. TO RESECTION MARGIN:

1 mm to dorsal margin

- NO. OF LYMPH NODES EXAMINED: 13
- NO. OF LYMPH NODES WITH TUMOR: 5
- HORMONE RECEPTOR STATUS:
 - Positive for estrogen receptor
 - Negative for progesteron receptor
- Her-2 STATUS:

Negative (2+ by immunohistochemistry, negative by FISH)

- pTNM CLASSIFICATION: pT1N2Mx
- EXTRA-TUMORAL BREAST: Fibrocystic disease

The Pathology report MAIN DIAGNOSIS:

 Local resection (lumpectomy) of breast with invasive ductal carcinoma

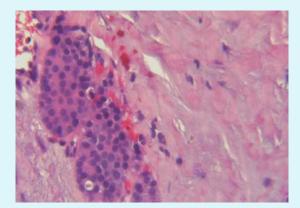
OTHER FEATURES:

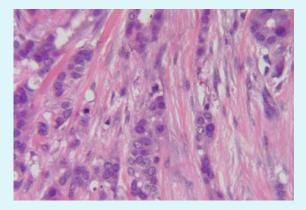
- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast
- TUMOR GRADE: 2
- <u>RESECTION MARGINS, MINIMUM DISTANCE TO TUMOR</u>: 4 mm from dorsal margin, 32 mm from ventral margin
- <u>TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE</u>: Absent.
- VASCULAR INVASION: Present
- DCIS: Present

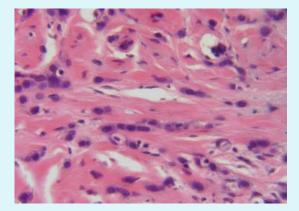
DCIS GRADE: 3 DCIS MIN. DIST. TO RESECTION MARGIN: 1 mm to dorsal margin

- NO. OF LYMPH NODES EXAMINED: 13
- NO. OF LYMPH NODES WITH TUMOR: 5
- HORMONE RECEPTOR STATUS:
 Positive for estrogen receptor
 Negative for progesteron receptor
- Her-2 STATUS:
 - Negative (2+ by immunohistochemistry, negative by FISH)
- pTNM CLASSIFICATION: pT1N2Mx
- EXTRA-TUMORAL BREAST: Fibrocystic disease

Types of invasive carcinoma



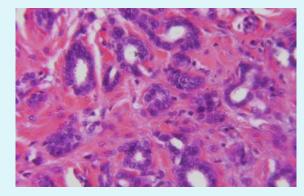


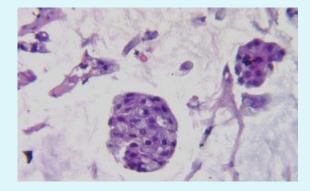


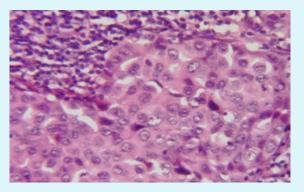
Ductal

Ductal

Lobular







Tubular

Mucinous

Medullary

Inflammatory carcinoma

• The diagnosis is <u>based on clinical features</u>:

Diffuse erythema, peau d'orange, tenderness, induration, warmth, enlargement

• And <u>carcinoma confirmed by biopsy:</u>

In most cases an invasive ductal carcinoma grade 3 with tumor in dermal lymphatics

MAIN DIAGNOSIS:

• Local resection (lumpectomy) of breast with invasive ductal carcinoma

OTHER FEATURES:

- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast

• TUMOR GRADE (1-3): 2

- <u>RESECTION MARGINS, MINIMUM DISTANCE TO TUMOR</u>: 4 mm from dorsal margin, 32 mm from ventral margin
- <u>TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE</u>: Absent.
- VASCULAR INVASION: Present
- DCIS: Present

DCIS GRADE: 3

DCIS MIN. DIST. TO RESECTION MARGIN:

1 mm to dorsal margin

- NO. OF LYMPH NODES EXAMINED: 13
- NO. OF LYMPH NODES WITH TUMOR: 5
- HORMONE RECEPTOR STATUS:
 Positive for estrogen receptor
 Negative for progesteron receptor
- Her-2 STATUS:

Negative (2+ by immunohistochemistry, negative by FISH)

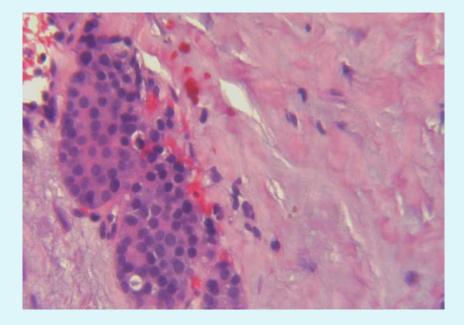
- pTNM CLASSIFICATION: pT1N2Mx
- EXTRA-TUMORAL BREAST: Fibrocystic disease

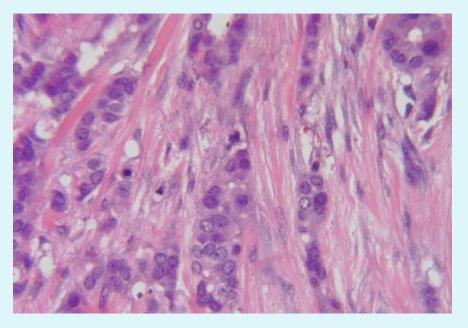
Tumor grading

Feature	Score
Tubule formation	1-3
Nuclear atypia	1-3
Number of mitoses	1-3

Grade	Sum of scores
Grade 1	3-5
Grade 2	6-7
Grade 3	8-9

Tumor grades





Grade 1

Grade 3

MAIN DIAGNOSIS:

Local resection (lumpectomy) of breast with invasive ductal carcinoma

OTHER FEATURES:

- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast
- TUMOR GRADE: 2

<u>RESECTION MARGINS, MINIMUM DISTANCE</u> <u>TO TUMOR</u>: 4 mm from dorsal margin, 32 mm from ventral margin

- <u>TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE</u>: Absent.
- VASCULAR INVASION: Present
- DCIS: Present

DCIS GRADE: 3 DCIS MIN. DIST. TO RESECTION MARGIN:

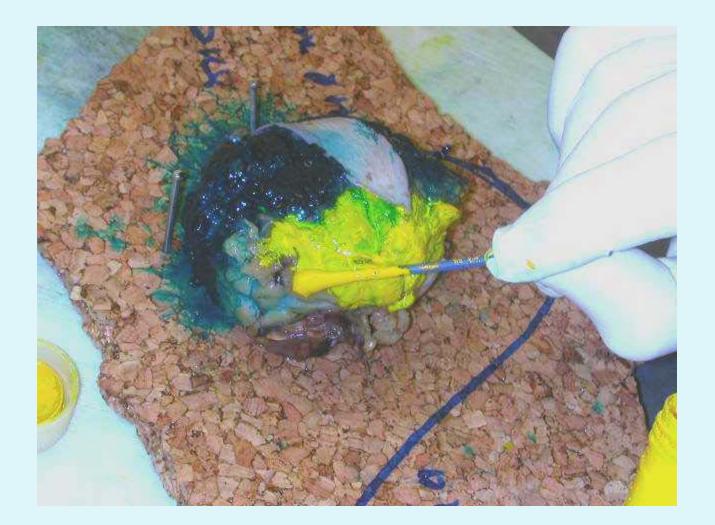
1 mm to dorsal margin

- NO. OF LYMPH NODES EXAMINED: 13
- NO. OF LYMPH NODES WITH TUMOR: 5
- HORMONE RECEPTOR STATUS: Positive for estrogen receptor Negative for progesteron receptor
- Her-2 STATUS:
 - Negative (2+ by immunohistochemistry, negative by FISH)
- pTNM CLASSIFICATION: pT1N2Mx
- EXTRA-TUMORAL BREAST: Fibrocystic disease

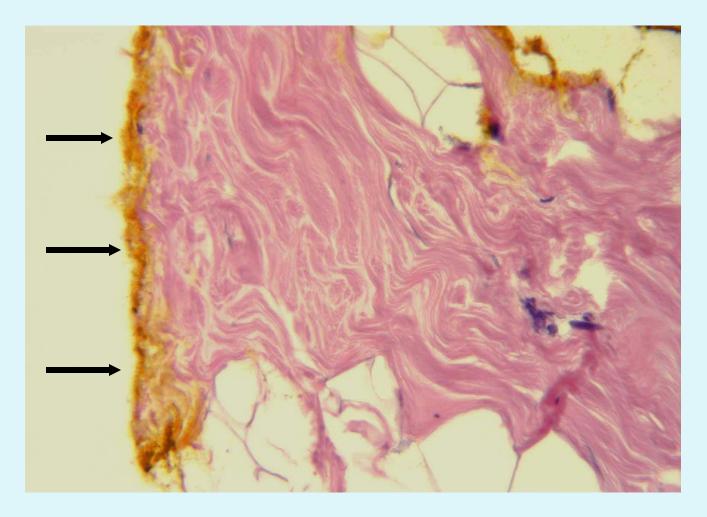
Orientation of the specimen



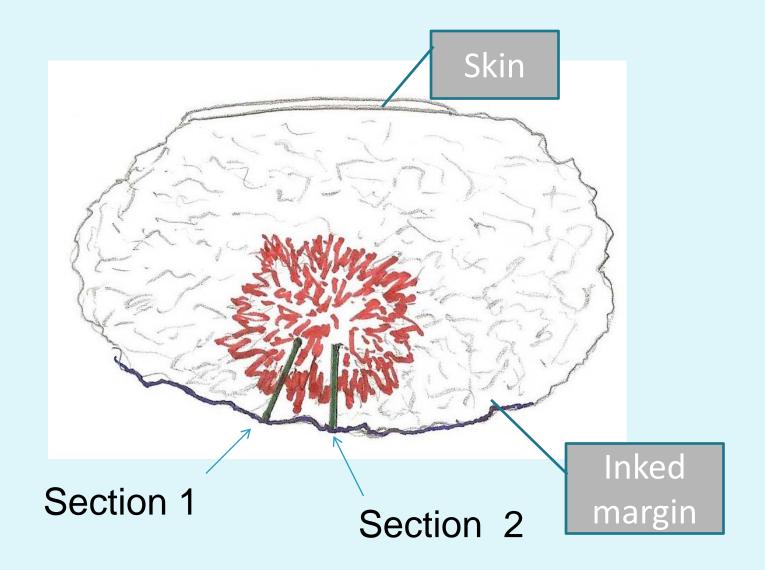
Inking of resection margins



The inked surface as seen in the microscopic slide



Examination of resection margin



MAIN DIAGNOSIS:

• Local resection (lumpectomy) of breast with invasive ductal carcinoma

OTHER FEATURES:

- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast
- TUMOR GRADE: 2
- <u>RESECTION MARGINS, MINIMUM DISTANCE TO TUMOR</u>: 4 mm from dorsal margin, 32 mm from ventral margin
- <u>TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE</u>: Absent.

VASCULAR INVASION: Present

DCIS: Present

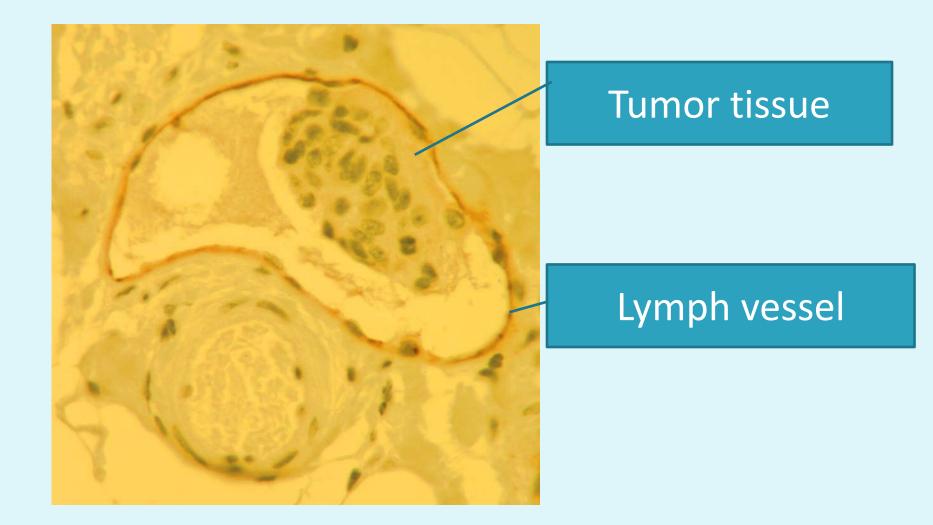
DCIS GRADE: 3 DCIS MIN. DIST. TO RESECTION MARGIN: 1 mm to dorsal margin

- NO. OF LYMPH NODES EXAMINED: 13
- NO. OF LYMPH NODES WITH TUMOR: 5
- HORMONE RECEPTOR STATUS: Positive for estrogen receptor Negative for progesteron receptor
- Her-2 STATUS:

Negative (2+ by immunohistochemistry, negative by FISH)

- pTNM CLASSIFICATION: pT1N2Mx
- EXTRA-TUMORAL BREAST: Fibrocystic disease

Vascular invasion



MAIN DIAGNOSIS:

Local resection (lumpectomy) of breast with invasive ductal carcinoma

OTHER FEATURES:

- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast
- TUMOR GRADE: 2
- <u>RESECTION MARGINS, MINIMUM DISTANCE TO TUMOR</u>: 4 mm from dorsal margin, 32 mm from ventral margin
- TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE: Absent.
- VASCULAR INVASION: Present

<u>DCIS</u>: Present DCIS GRADE (1-3): 3 DCIS MINIMUM DISTANCE TO RESECTION MARGIN: 1 mm to dorsal margin

- NO. OF LYMPH NODES EXAMINED: 13
- NO. OF LYMPH NODES WITH TUMOR: 5
- HORMONE RECEPTOR STATUS: Positive for estrogen receptor Negative for progesteron receptor
- Her-2 STATUS: Negative (2+ by immunohistochemistry, negative by FISH)
- pTNM CLASSIFICATION: pT1N2Mx
- EXTRA-TUMORAL BREAST: Fibrocystic disease

MAIN DIAGNOSIS:

• Local resection (lumpectomy) of breast with invasive ductal carcinoma

OTHER FEATURES:

- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast
- TUMOR GRADE: 2
- <u>RESECTION MARGINS, MINIMUM DISTANCE TO TUMOR</u>: 4 mm from dorsal margin, 32 mm from ventral margin
- <u>TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE</u>: Absent.
- VASCULAR INVASION: Present
- DCIS: Present

DCIS GRADE: 3 DCIS MIN. DIST. TO RESECTION MARGIN: 1 mm to dorsal margin

• NO. OF LYMPH NODES EXAMINED: 13

- NO. OF LYMPH NODES WITH TUMOR: 5
- HORMONE RECEPTOR STATUS:

Positive for estrogen receptor Negative for progesteron receptor

• Her-2 STATUS:

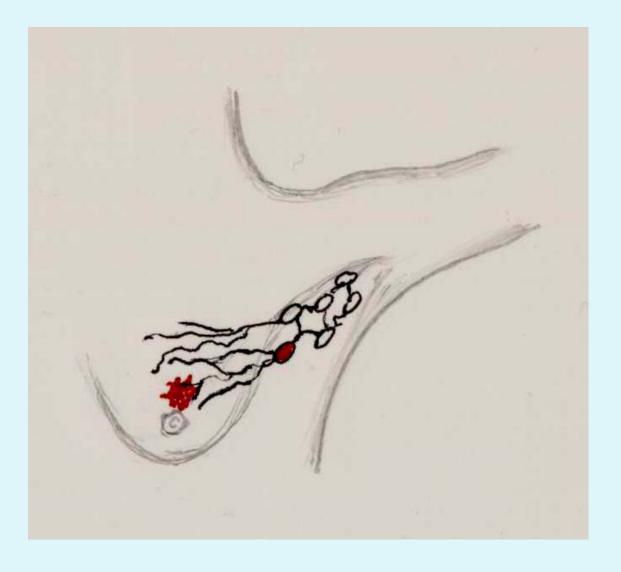
Negative (2+ by immunohistochemistry, negative by FISH)

- pTNM CLASSIFICATION: pT1N2Mx
- EXTRA-TUMORAL BREAST: Fibrocystic disease

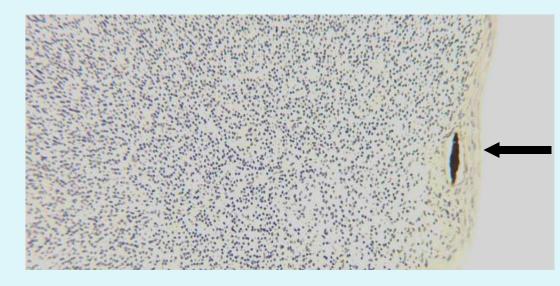
Axillary lymph nodes

Breast cancer spreads through lymphatic channels to axillary lymph nodes.

When axillary content is removed, all nodes are searched and embedded for microscopy



Micrometastasis



Micrometastasis spotted in otherwise negative sentinel node

Tumor deposits in lymph node

Size	Designated	pTNM class
> 2 mm	Metastasis	pN1
≤ 2 mm; > 0.2 mm	Micrometastasis	pN1(mi)
≤ 0.2 mm	Isolated tumor cells	pN0 (i+)

MAIN DIAGNOSIS:

• Local resection (lumpectomy) of breast with invasive ductal carcinoma

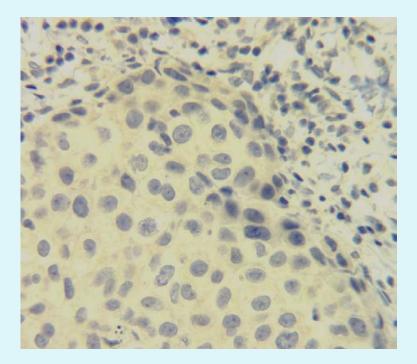
OTHER FEATURES:

- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast
- TUMOR GRADE: 2
- <u>RESECTION MARGINS, MINIMUM DISTANCE TO TUMOR</u>: 4 mm from dorsal margin, 32 mm from ventral margin
- <u>TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE</u>: Absent.
- VASCULAR INVASION: Present
- DCIS: Present
 - DCIS GRADE: 3 DCIS MIN. DIST. TO RESECTION MARGIN:
 - 1 mm to dorsal margin
- NO. OF LYMPH NODES EXAMINED: 13
- NO. OF LYMPH NODES WITH TUMOR: 5

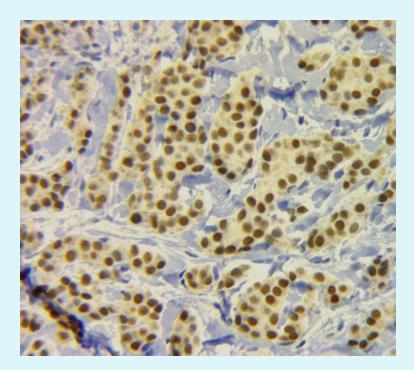
HORMONE RECEPTOR STATUS: Positive for estrogen receptor Negative for progesteron receptor

- Her-2 STATUS:
 - Negative (2+ by immunohistochemistry, negative by FISH)
- pTNM CLASSIFICATION: pT1N2Mx
- EXTRA-TUMORAL BREAST: Fibrocystic disease

Immunohistochemistry: Estrogen receptor



Negative: No benefit from hormone therapy



Positive: Will benefit from hormone therapy

MAIN DIAGNOSIS:

• Local resection (lumpectomy) of breast with invasive ductal carcinoma

OTHER FEATURES:

- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast
- TUMOR GRADE: 2
- <u>RESECTION MARGINS, MINIMUM DISTANCE TO TUMOR</u>: 4 mm from dorsal margin, 32 mm from ventral margin
- <u>TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE</u>: Absent.
- <u>VASCULAR INVASION</u>: Present
- DCIS: Present

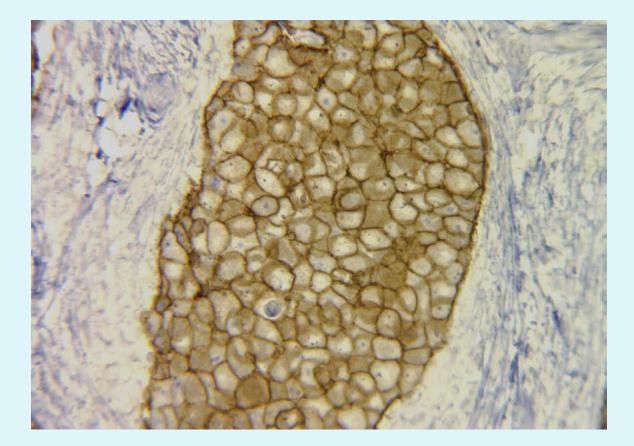
DCIS GRADE: 3 DCIS MIN. DIST. TO RESECTION MARGIN: 1 mm to dorsal margin

- NO. OF LYMPH NODES EXAMINED: 13
- NO. OF LYMPH NODES WITH TUMOR: 5
- HORMONE RECEPTOR STATUS:
 Positive for estrogen receptor
 Negative for progesteron receptor

• Her-2 STATUS: Negative

- pTNM CLASSIFICATION: pT1N2Mx
- EXTRA-TUMORAL BREAST: Fibrocystic disease

Immunohistochemistry Her-2



Her-2 positive. Will benefit from Herceptin therapy

MAIN DIAGNOSIS:

• Local resection (lumpectomy) of breast with invasive ductal carcinoma

OTHER FEATURES:

- TUMOR SIZE: 3 cm
- LOCATION: Upper inner quadrant, right breast
- TUMOR GRADE: 2
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- <u>TUMOR INVASION OF SKIN, MUSCLE, PAGET'S DISEASE OF THE NIPPLE</u>: Absent.
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- Her-2 STATUS:

Negative (2+ by immunohistochemistry, negative by FISH)

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• EXTRA-TUMORAL BREAST: Fibrocystic disease

MAIN DIAGNOSIS:

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Negative (2+ by immunohistochemistry, negative by FISH)

• pTNM CLASSIFICATION: pT1N2Mx

• EXTRA-TUMORAL BREAST: Fibrocystic disease



Is Breast pathology in Ghana similar to that in the industralized countries

A comparison of non-inflammatory pathology diagnoses in Ghana (KATH) and Norway (UNN)

The relative frequency of major pathology diagnoses in breast specimens from KATH and UNN 2006-2009

Diagnosis	Ghana, KATH (162)		Norwa (21	y, UNN 73)
	Ν	%	Ν	%
Fibroadenoma	46	28 %	311	14 %
Fibrocystic dis.	20	12 %	524	24 %
Atypical hyperpl	0	0 %	28	1 %
Carcinoma in situ	6	4 %	301	14 %
Carcinoma	90	56 %	1009	46 %

The relative frequency of main types of invasive breast carcinoma 2006-2009

Туре	Ghana, KATH (92)			y, UNN 12)
	Ν	%	Ν	%
Ductal and NOS	85	92,4	873	86,3
Lobular	1	1,1	97	9,6
Tubular	0	0,0	10	1,0
Mucinous	3	3,3	25	2,5
Medullary	1	1,1	3	0,3
Papillary	2	2,2	3	0,3

Tumor grade in Core biopsies of Breast cancer Preliminary data 2009

Tumor grade	Ghana (25 cases)	Norway (93 cases)
Grade 1	16%	33%
Grade 2	28%	44%
Grade 3	48%	16%

Hormone receptors in Core biopsies of Breast cancer Preliminary data 2009

Hormone receptor	Ghana (25 cases)	Norway (93 cases)
Estrogen receptor	48% positive	89% positive
Progesteron receptor	12% positive	62% positive

Her-2 overexpression in Core biopsies of Breast cancer Preliminary data 2009

Ghana	Norway
(25 cases)	(93 cases)
32% positive	12% positive