UUID:CFF567FC-8867-4CA4-85AE-81A7D30095E7 TCGA-A1-A0SE-01A-PR Redacted

105-0-3 Carcinoma, nigiltrating lubular and ductal 8522/3 Site: bread, Nos C50.9 M 10/21/11

Final Pathologic Diagnosis:

- A. Right breast, biopsy: No evidence of tumor in one lymph node (0/1).
- B. Left axilla, sentinel lymph node #1, dissection: No evidence of tumor in three lymph nodes (0/3).
- C. Right breast, mastectomy:
- 1. Skin with no significant pathologic abnormality.
- 2. Proliferative fibrocystic changes.
- 3. Microcalcifications in association with benign glands.
- D. Left breast, mastectomy:
- 1. Invasive carcinoma, 2.0 cm, SBR grade 2, see comment.
- 2. Ductal carcinoma in situ, low grade, see comment.
- 3. Skin with no significant pathologic abnormality.
- 4. Fibroadenoma.
- 5. Microcalcifications associated with invasive carcinoma and benign glands.
- 6. Proliferative fibrocystic changes.

- 7. Papillomatosis.
- 8. Pseudoangiomatous stromal hyperplasia.
- E. Nonsentinel lymph node, left axilla, dissection: No evidence of tumor in four lymph nodes (0/4).
- F. Right breast, re-excision of superior portion: Benign breast parenchyma. Note: In Part D (left breast), the area indicated by the surgeon with a green stitch shows proliferative fibrocystic changes, sclerosing adenosis, and pseudoangiomatous stromal hyperplasia. There is no evidence of carcinoma in the area around the green suture.

Breast Tumor Synoptic Comment

- Laterality: Left.
- Invasive tumor type: Mixed ductal and lobular type (tumor is present in Slides D1, D2, D5, and D18).
- Invasive tumor size: 2.0 cm maximum diameter (Slide D5).
- Invasive tumor grade (modified Bloom-Richardson):

Nuclear grade: 2 points.

Mitotic count: 16 mitotic figures/10 HPF, 2 points.

Tubule/papilla formation: Definite tubule formation <10%, 3 points.

Total points and SBR grade = 7 points, grade 2.

- Lymphatic-vascular invasion: None identified.
- Perineural invasion: None identified.
- Resection margins for invasive tumor:
 - Deep margin: Positive (Slide D2, 5).
 - Medial margin: Widely clear; >1 cm.
 - Lateral margin: Widely clear; >1 cm.
 - Anterior/superior margin: Widely clear; >1 cm.
 - Anterior/inferior margin: Widely clear; >1 cm.
- Ductal carcinoma in situ (DCIS) type: Solid.
- Ductal carcinoma in situ size: DCIS present as scattered microscopic foci associated with the invasive component on Slides D1 and D2.
- Ductal carcinoma in situ nuclear grade: Low grade.
- Necrosis in ductal carcinoma in situ: None.
- Microcalcifications: Present involving invasive carcinoma.
- Resection margins for ductal carcinoma in situ:
 - Deep margin: Close; within less than 0.3 cm (Slide D1).
 - Medial margin: Widely clear: >1 cm.
 - Lateral margin: Widely clear; >1 cm.
 - Anterior/superior margin: Widely clear; >1 cm.
 - Anterior/inferior margin: Widely clear; >1 cm.
- Lobular carcinoma in situ (LCIS): Not identified.
- Lymph node status: No evidence of tumor in eight lymph nodes (0/8).
- AJCC/UICC stage: pT1cN0MX.
- Nontumorous breast tissue: Proliferative fibrocystic changes, Fibroadenoma, pseudoangiomatous hyperplasia.
- Nipple: Unremarkable.
- Skin/dermis: Unremarkable.

Immunohistochemical tests for estrogen, progesterone and Her2 Neu are pending and will follow in an addendum.

Intraoperative Consult Diagnosis

FS1 (A) Right breast nodule, biopsy: Lymph node with no tumor seen. (Dr.

FS2 (B) Sentinel lymph node, left axilla, biopsy: Three lymph nodes, no tumor seen. (Dr.



Clinical History

The patient is a year-old woman with breast cancer. A tracking sheet accompanies the specimen and indicates a high family risk for cancer and preoperative diagnosis of known left breast cancer in the upper outer quadrant. No specimen radiograph is requested. Multifocality is suspected on the left side. Special stains for ER, PR, and HER2/neu are requested as a repeat. The diagram indicates that the known tumor is in the upper outer quadrant of the left breast, and a suspicious area is present in the upper inner quadrant at approximately the 11 o'clock position above the nipple. Recent MRI indicates the presence of a known 2-cm tumor with clip present in the lateral breast of the left breast in addition to an adjacent 5-mm lesion that is immediately posteromedial to the primary lesion. A 5-mm suspicious area is also identified in the slightly upper inner quadrant.

Gross Description

The specimen is received in six parts, each labeled with the patient's name and medical record number. Parts A through D and F are received fresh, and Part E is received in formalin.

Part A is additionally labeled "right breast nodule." It consists of a single unoriented, irregular piece of soft, pink tissue, measuring 0.7 x 0.5 x 0.3 cm. The specimen is entirely submitted for frozen section diagnosis as FS1, with the frozen section remnant submitted in cassette A1.

Part B is additionally labeled "sentinel lymph node left axilla, count = 1800, frozen section." It consists of a single irregular piece of soft, yellow-pink, fatty tissue, measuring 3.0 x 2.0 x 0.6 cm. The specimen is entirely submitted for frozen section diagnosis as FS2, with the frozen section remnant submitted in cassette B1.

Part C is additionally labeled "right breast." It consists of a mastectomy specimen, oriented with a short suture considered superior and long suture considered lateral; oriented as such, the specimen measures 3.2 cm from anterior to posterior, 14.7 cm from medial to lateral, and 13.7 cm from superior to inferior. The mastectomy specimen weighs 212.5 gm. The specimen includes an area of skin, measuring 14.5 x 6.7 cm, with nipple, measuring 1.7 x 1.7 x 1.5 cm, and areola. measuring 3 x 3.8 cm. The specimen has been previously inked and serially sectioned by the slices. Inking follows standard inking, with posterior black, anterior superior blue, and anterior inferior green. The accompanying research paperwork indicates that a fragment of tissue has been taken for tissue banking. Cut sections reveal that the breast is composed almost entirely of homogeneous, firm, white, fibrous tissue that abuts the deep margin. Thin layers of adipose tissue are present in the anterior surface as well as the medial aspect of the specimen. No grossly evident lesions are identified. The nipple and skin similarly appear unremarkable. Slice 1 is considered medial, and slice 10 is considered lateral. The nipple, thereby, is in slice 5. Representative sections are submitted as follows:

Cassettes C1-C2: Nipple, entirely submitted. Cassette C3: Upper outer quadrant from slice 8. Cassette C4: Upper outer quadrant from slice 7. Cassette C5: Lower outer quadrant from slice 8. Cassette C6: Lower outer quadrant from slice 7. Cassette C7: Upper inner quadrant from slice 4. Cassette C8: Upper inner quadrant from slice 3. Cassette C9: Lower inner quadrant from slice 3. Cassette C10: Lower inner quadrant from slice 2 and section of skin from slice 4.

Part D is additionally labeled "left breast," with further specification in the requisition form indicating that a green stitch = questionable secondary cancer, correlate with MRI, and black short = superior, long = lateral.

A firm, pale-tan, circumscribed lesion, measuring 1.6 x 1.2 x 1.4 cm, is present at the 3 o'clock position, 3 cm from the nipple; this contains a surgical clip. The lesion is 0.1 cm from the deep margin, 0.9 cm from the skin, 2.2 cm from anterior superior, 1.9 cm from anterior inferior, 3.5 cm from the lateral margin, and <10 cm from the medial margin. Immediately inferior and medial to main lesion, a 0.4-cm area contains multiple punctate, firm, yellow spots. This lesion is 0.5 cm from the large lesion and is 0.5 from the deep margin. The remainder of the specimen is composed mostly of firm, fibrous tissue, showing multiple small cystic areas, with a maximum diameter of 0.4 cm, throughout. The area adjacent to the green suture on slice 6 similarly consists of dense, fibrous tissue, and no definitive lesions are identified in this area. The skin, nipple, and areola appear unremarkable.



Surgical Pathology - Working Draft Accompanying paperwork from the ' ndicates that a portion of the specimen has been taken for tissue banking. The specimen contains a short stitch and long black stitch, taken to be superior and lateral, respectively, and has been previously inked and sectioned by the research technician into ten slices, from medial to lateral. Inking is as per standard, with posterior in black, anterior superior in blue, and anterior inferior in green. The blue ink appears faint. The most-medial slice is taken as slice 1 and the most-lateral slice as slice 10. The specimen, thereby, measures 2.5 cm from anterior to posterior, 15.6 cm from medial to lateral, and 16.6 cm from superior to inferior and weighs 192.5 gm. A skin ellipse is present, measuring 6.6 x 13.1 cm, with the long axis in the mediolateral extent. The areola measures 3.4 x 3 cm, and the nipple measures 1.8 x 1.6 x 1.6 cm. The nipple lies in slices 5 and 6. The green nylon suture is in the posterior aspect of slice 6 at the 12 o'clock position, 3.5 cm superior to the nipple. Representative sections are submitted as follows: Cassette D1: 1.6-cm lateral lesion in relation to deep margin from slice 8. Cassette D2: Punctate, yellow lesion from slice 8. Cassette D3: Nearest anterior-superior margin in slice 8. Cassette D4:

Nearest anterior-inferior margin from slice 8.

Cassette D5:

Additional 1.6-cm lesion in relation to skin.

Cassettes D6-D7:

Nipple, entirely submitted.

Cassette D8:

Area marked by green suture, including skin and anterior-superior and deep

margins.

Cassettes D9-D11:

Additional sections of area marked by the green suture en bloc from slice 6.

Cassette D12:

Area adjacent to green suture from slice 5. Area adjacent to green suture from slice 7.

Cassette D13: Cassette D14:

Inferior margin of slice 8. Superior margin of slice 8.

Cassette D15: Cassette D16:

Lateral margin from slice 10. Medial margin from slice 1.

Cassette D17: Cassette D18:

Intervening area between primary lesion and lateral margin from slice 9.

Cassette D19: Cassette D20:

Upper inner quadrant, rectangular from slice 4 and triangular from slice 3. Lower inner quadrant, rectangular from slice 5, triangular from slice 4.

Cassette D21: Cassette D22:

Upper outer quadrant, rectangular from slice 8, triangular from slice 9. Lower outer quadrant, triangular from slice 7, rectangular from slice 9.

Part E is additionally labeled "nonsentinel lymph node left axilla." It consists of a single unoriented fragment of largely adipose tissue, measuring 2.4 x 1.4 x 0.5 cm. A single 1.4-cm lymph node is identified. The candidate lymph node is bisected and entirely submitted in cassette E1. The remainder of the soft tissue is entirely submitted in cassette E2.

Part F is additionally labeled "re-excision superior portion right breast" and has an additional note in the requisition form indicating that the stitch = new margin. It consists of a single fragment of adipose tissue, measuring 3 x 2.2 x 0.9 cm, marked by a single black suture. The surface containing the black suture is inked in black and the opposite surface in blue. The specimen is serially sectioned to reveal mostly adipose tissue, with a single 0.7-cm area of white, fibrous tissue. The specimen is entirely submitted in cassettes F1 through F3.

/Pathology Resident

■/Pathologist Signed:

Fee Codes:

Addenda

Addendum.

Date Ordered:

Date Complete:

Date Reported:

Status: Signed Out



Addendum Comment

An immunohistochemical test for estrogen and progesterone receptors as well as for HER2 was performed on block D5.

The test for estrogen receptors is positive. There is strong (3+) nuclear staining in 85% of tumor cells. Internal positive control is positive.

The test for progesterone receptors is. There is strong (3+) nuclear staining in~100% of tumor cells. Internal positive control is positive.

Result of HER2/neu test: This carcinoma is negative for HER2/neu oncoprotein over-expression.

An immunohistochemical assay was performed using the CB11 monoclonal antibody to HER2/neu oncoprotein. The staining intensity of this carcinoma was 1 on a scale of 0-3 (HER2 test interpreted by Dr.

Carcinomas with staining intensity scores of 0 or 1 are considered negative for over-expression of HER2/neu oncoprotein.

Those with a staining intensity score of 2 are considered borderline. We and others have observed that many carcinomas with staining intensity scores of 2 do not show gene amplification. All carcinomas with staining intensity scores of 2 are therefore submitted for FISH testing. The results of the FISH test are issued directly from the molecular cytogenetics laboratory.

Carcinomas with staining intensity scores of 3 are considered positive for over-expression of HER2/neu oncoprotein. Tumors in this category show an excellent correlation between the results of immunohistochemical and FISH testing, and almost always show gene amplification.

The immunoperoxidase stain(s) reported above were developed and their performance characteristics determined by the They have not been cleared or approved by the U. S. Food and Drug Administration. Ine FDA has determined that such clearance or approval is not necessary. These tests are used for clinical purposes. They should not be regarded as investigational or for research. This laboratory is certified under the Clinical Laboratory Improvement Amendments of 1988 ("CLIA") as qualified to perform high-complexity clinical testing.

> ■ Pathologist Electronically signed out on

Addendum.

Date Ordered: Date Complete: Date Reported:

Status: Signed Out

Addendum Comment

An immunohistochemical test for progesterone receptors was performed on block D5.

The test for progesterone receptors is positive. There is strong nuclear staining in ~100% of tumor cells.

The immunoperoxidase stain(s) reported above were developed and their performance characteristics determined by the They have not been cleared or approved by the U. S. Food and Drug Administration. The FDA nas determined that such clearance or approval is not necessary. These tests are used for clinical purposes. They should not be regarded as investigational or for research. This laboratory is certified under the Clinical Laboratory Improvement Amendments of 1988 ("CLIA") as qualified to perform high-complexity clinical testing.

> Pathologist Electronically signed out on



Surgical I	Pathology - Worki	ing Draft
SPECIMEN ADEQUACY:		
Satisfactory for even endocervical/trans	aluation; atrophic patte formation zone compor	ern with no identifiable nent.
Specimen Class:	Status: Signed Out	Accessioned:
Specimen(s) Received: Cervical/Endocervica Final Diagnosis	l, Thin Prep Imaged	Signed Out:
Cervical/Endocervical, Thin Pre	ep Imaged	
NEGATIVE FOR INTRAEP. Atrophic changes	ITHELIAL LESION OR M	MALIGNANCY.
SPECIMEN ADEQUACY: Satisfactory for eva endocervical/transf	aluation; atrophic patte formation zone compor	ern with no identifiable nent.
Specimen Class:	Status: Signed Out	Accessioned Signed Out:
Specimen(s) Received: Left breast core needle Final Diagnosis		-
Left breast, needle core biopsy:	Invasive carcinoma; see	comment.
Procedure/Addends for ADDENDUM.	Date of Addendum.	:
staining. Thus, this invasive carcinon	staining and others that are na displays features of both mohistochemical stained sliced above were developed an Administration. The FDA has are used for clinical purposaboratory is certified under the staining and staining the staining and staining an	ductal and lobular carcinoma. Dr. de and concurs. In their performance characteristics They have not been cleared or as determined that such clearance or ses. They should not be regarded as the Clinical Laboratory Improvement

Status: Signed Out

Accessioned: Signed Out:

Specimen Class:

Specimen(s) Received: Left Axilla Lymph Node, Fine Needle Aspiration

			ì		
Surgical Pathology - Working Draft					
	Final Diagnosis Left Axilla Lymph Node, Fine Needle Aspiration: Benign reactive lymph node, see note.				
			· -		
This lymph tingible boo	node shows a spectrum ly macrophages. No ev	m of small to large si ridence of neoplasia i	zed lymphocytes with scattered is seen.		
	_Specimen Class:	Status: Signed Out	Accessioned: Signed Out:		
	eceived: Left Breast, Fine Needle	Aspiration	Signed Out.		
Final Diagnosis Left Breast	, Fine Needle Aspiration	n: Adenocarcinoma	, see comment.		
Procedure/Add	enda for				
ADDENDU	IM.	Date of Addendum	1.:		
Addendum Comment An immunohistochemical test for estrogen and progesterone receptors as well as for HEr-2-neu was performed on the material submitted for cell block.					
The test for estrogen receptors is positive. There is strong nuclear staining in 90% of tumor cells.					
The test for progesterone receptors is positive. There is strong nuclear staining in 80% of tumor cells.					
Result of HER2/neu test: This carcinoma is borderline for HER2/neu oncoprotein over-expression.					
An immunohistochemical assay was performed on cell block using the CB11 monoclonal antibody to HER2/neu oncoprotein. The staining intensity of this carcinoma was 2 on a scale of 0-3.					
Carcinomas v HER2/neu or		es of 0 or 1 are considere	ed negative for over-expression of		
many carcino staining inter	mas with staining intensity	/ scores of 2 do not show re submitted for FISH te	ine. We and others have observed that we gene amplification. All carcinomas with esting. The results of the FISH test are		
oncoprotein.	with staining intensity score Tumors in this category sh chemical and FISH testing,	ow an excellent correlat	ositive for over-expression of HER2/neution between the results of w gene amplification.		
Dr	nas reviewed the HER2/neu	stain and concurs.			