AMSER Case of the Month March 2021

46-year-old female with left breast mass



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Patient Presentation

- HPI: 46-year-old female presents for an annual gynecological exam, offering complaints. No history of recent weight loss, fever or fatigue. Denies lumps in her beast, nipple discharge, or breast tenderness; no swollen lymph nodes in the groin or axilla.
- PSH: Supracervical hysterectomy, retaining both ovaries
- OB/GYN: No history of abnormal mammogram. As per patient, prior mammogram 1 year ago at OSH was normal (images unavailable for review).
- PE: Overall symmetric breasts. Large, firm golf ball size lump palpated in the left breast at the 12 o-clock. No associated skin changes.



What imaging should we order?



ACR Appropriateness Criteria: Palpable Breast Masses

This imaging modality was ordered. Variant 1:

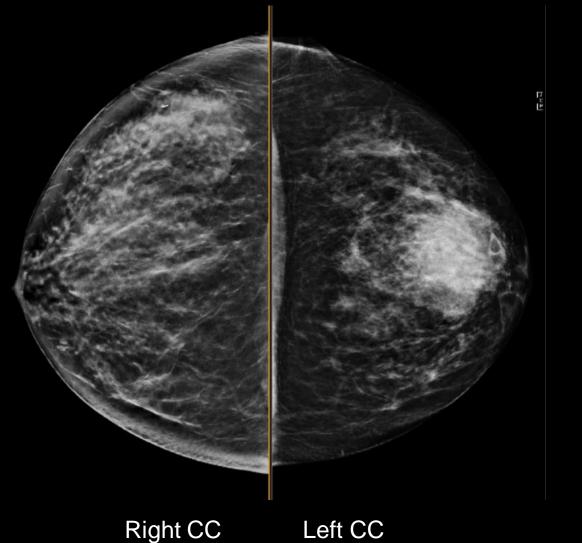
IA-IB for additional steps in the workup of these patients.)				
Radiologic Procedure	Rating	Comments	RRL*	
Mammography diagnostic	9	See references [13-15].	**	
Digital breast tomosynthesis diagnostic	9	See references [16-18,20,85].	**	
US breast	4	If she had recent mammogram (ie, past 6 months), US may be appropriate.	0	
MRI breast without and with IV contrast	2	See references [4,49].	0	
MRI breast without IV contrast	1		0	
FDG-PEM	1		€€€	
Sestamibi MBI	1		***	
Image-guided core biopsy breast	1		Varies	
Image-guided fine-needle aspiration breast	1		Varies	
Rating Scale: 1,2,3 Usually not appropriate; 4,5,6 May be appropriate; 7,8,9 Usually appropriate			*Relative Radiation Level	

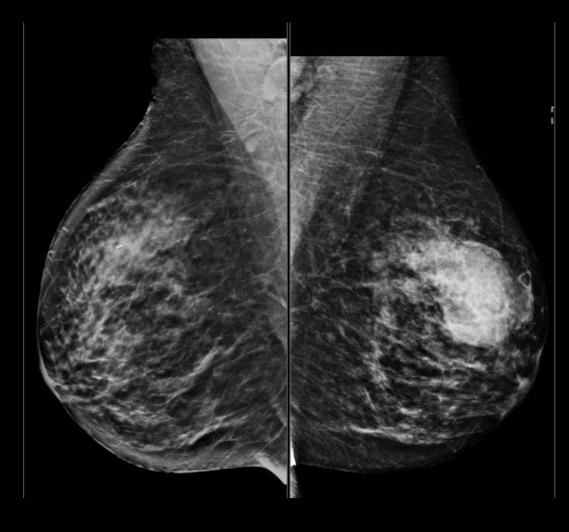
 1_{A-1B} for additional steps in the workup of these patients)

Palpable breast mass. Female, 40 years of age or older, initial evaluation. (See Appendices

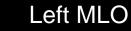


Diagnostic Mammogram (unlabeled)



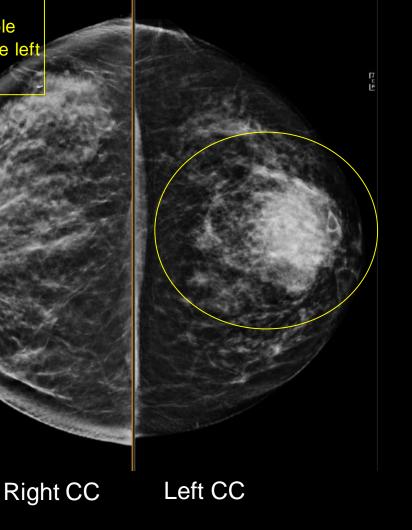


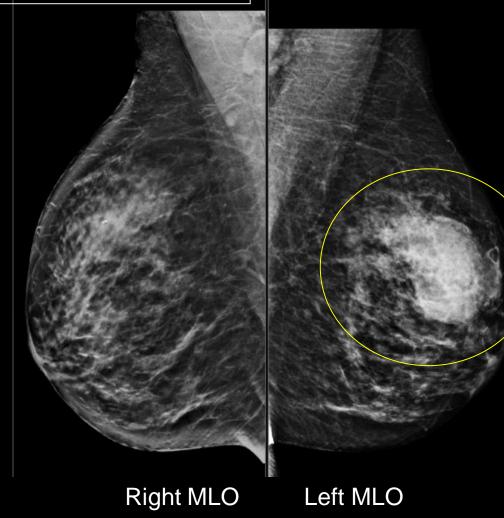
Right MLO



Diagnostic Mammogram Findings (labeled)

Heterogeneously dense breast tissue with a hyperdense mass with obscured borders underlying the palpable marker in 12 :00 in the left breast. There were no suspicious calcifications associated with the mass or any additional concerning findings in either breast.





What additional imaging should we order next?



ACR Appropriateness Criteria: Palpable Breast Masses

These imaging modalities were ordered. Variant 1:

<u>1A-1B</u> for additional steps in the workup of these patients.)				
Radiologic Procedure	Rating	Comments	RRL*	
Mammography diagnostic	9	See references [13-15].	**	
Digital breast tomosynthesis diagnostic	9	See references [16-18,20,85].	**	
US breast	4	If she had recent mammogram (ie, past 6 months), US may be appropriate.	0	
MRI breast without and with IV contrast	2	See references [4,49].	0	
MRI breast without IV contrast	1		0	
FDG-PEM	1		€€€€	
Sestamibi MBI	1		€€€	
Image-guided core biopsy breast	1		Varies	
Image-guided fine-needle aspiration breast	1		Varies	
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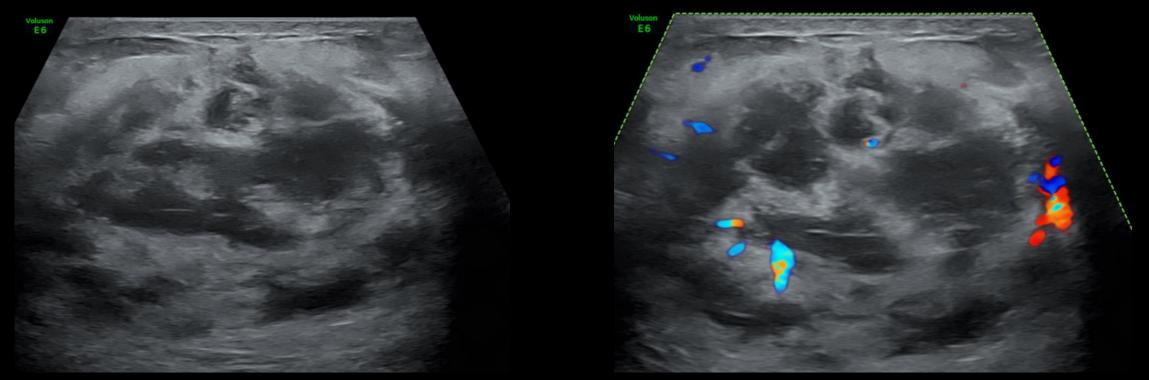
Palpable breast mass. Female, 40 years of age or older, initial evaluation. (See Appendices



Ultrasound of left breast mass

without Doppler

with Doppler

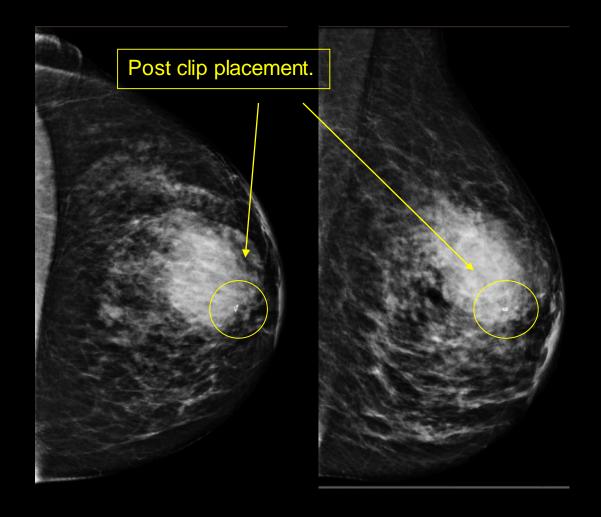


Sonographic evaluation of the left breast revealed a heterogeneous, solid and cystic 5.1 x 3.5 x 4.7 cm mass with indistinct margins and internal vascularity at 12:00, corresponding to mammogram finding. BIRADS 4. There was no left axillary adenopathy. Subsequent ultrasound guided core biopsy was performed.

Ultrasound Guided Core Biopsy & Post Biopsy During biopsy Clip Placement Post biopsy

14g needle from ultrasound guided biopsy device

Post clip placement



CC view

MLO view

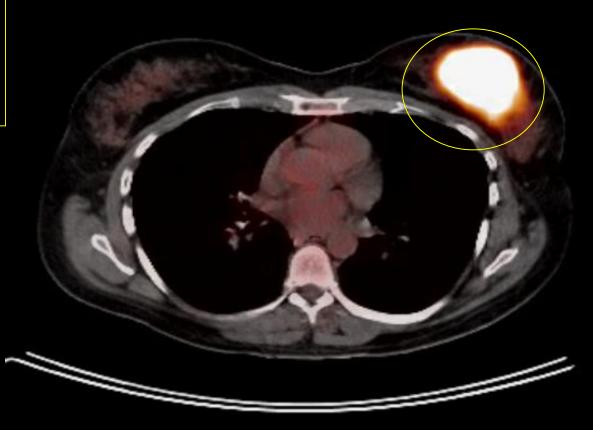
Findings and Final Dx:

- Pathology revealed a starry sky morphology and a phenotype of CD20+, CD79a+, CD10+, bcl6+, myc+, bcl2 and bcl1 negative, and TdT negative. Ki-67 was 99%. Fluorescence in situ hybridization revealed a bcl6 and c-myc rearrangement.
- The final pathology review classified the specimen as a double-hit high grade B cell lymphoma.



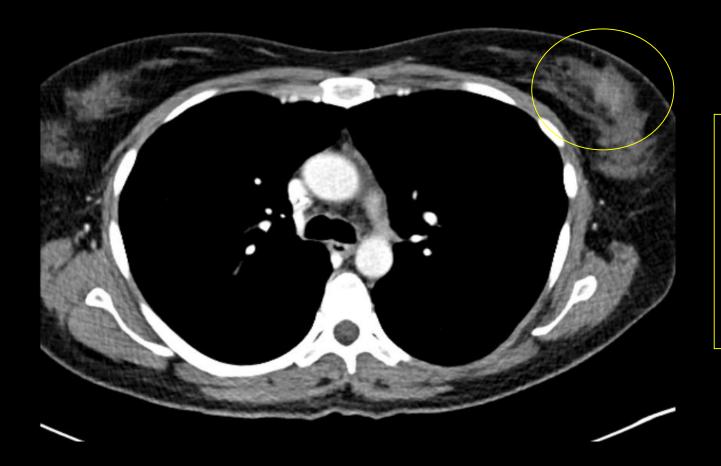
PET CT

Hypermetabolic 5.3 x 4.0 cm left breast mass, SUV max of 9.3



Approximately 1 month following initial diagnosis, a PET CT was performed.

CT CAP

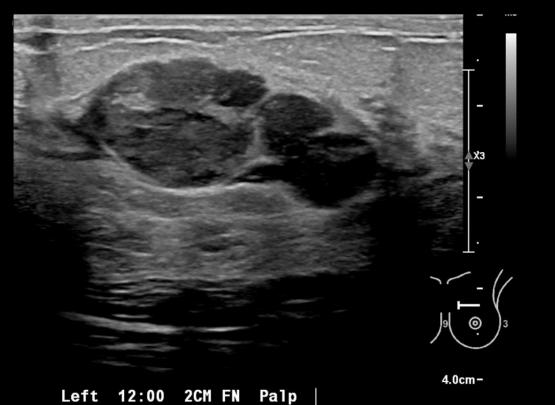


Ill-defined enhancing soft tissue in the left breast, decreased in size since original presentation; no distant metastases.

Restaging CT CAP was ordered 2 months following initial diagnosis.

3 month follow-up ultrasound

Without Doppler



With Doppler



Follow up ultrasound evaluation demonstrated a 3.3 x 1.6 x 2.1 cm hypoechoic mass at 12:00, 2 cm from nipple, a decrease in size from the original ultrasound imaging, with decreased vascularity and increasingly circumscribed margins; birads 6.

Final Imaging:

- Unfortunately, a repeat ultrasound performed 2 months after initial follow-up imaging (5 months since the original presentation) demonstrated an increase in tumor size, to 6.6 cm, initially 5.1 cm on presentation.
- Repeat ultrasound-guided biopsy revealed the same tumor as on initial biopsy, high-grade B-cell lymphoma.



Discussion: Primary Breast Lymphoma Presentation and Pathology

- Primary breast lymphoma is a form of extranodal lymphoma, representing 0.5% of all breast malignancies.
- Histologically, diffuse large B-cell lymphoma is the most common.
- PBL makes up 1% of all non-Hodgkin lymphoma and <3% of extranodal lymphomas.
- Most commonly presents solely as a painless breast mass, making it difficult to differentiate from breast carcinoma.
- Constitutional symptoms, cutaneous changes, and nipple retraction and discharge are not often associated.
- Right breast involvement is often reported over left breast.



Discussion: Imaging and Diagnosing

- While there are no distinguishing pathognomonic features on imaging, multiple studies denote features which could point to breast lymphoma over other pathologies.
- On mammography, circumscribed margins, absent calcifications, and normal breast architecture with axillary lymphadenopathy should propose lymphoma.
- On ultrasound, a hypoechoic mass either round or oval in nature is observed most commonly.
- PET/CT scans are beneficial in terms of staging and management.



Discussion: Treatment

- Treatment involves a combination of chemotherapy and radiotherapy. Additionally, CNS prophylaxis is also highly recommended.
- Suggested chemotherapy is R-CHOP (rituximab, cyclophosphamide, doxorubicin, vincristine, and prednisone).
- However, it has been proposed in cases of high Ki-67 expression and more aggressive lymphomas such as double- and triple-hit, R-EPOCH (etoposide, prednisone, vincristine, cyclophosphamide, and doxorubicin) is first-line.
 - This was the therapy chosen for the patient presented here with a Ki-67 of 99% diagnosed with double-hit high grade B cell lymphoma, in addition to intrathecal methotrexate and alternating cytarabine for CNS prophylaxis.



References:

- Cheah, C. Y., Campbell, B. A., & Seymour, J. F. (2014). Primary breast lymphoma. *Cancer treatment reviews*, 40(8), 900-908.
- Jia-Jia Huang, Wenqi Jiang, Zhi-Ming Li; R-EPOCH Is Superior to R-CHOP As a First-Line Regimen in De Novo DLBCL Patients with High Ki-67 Expression. *Blood* 2015; 126 (23): 5085. doi: <u>https://doi.org/10.1182/blood.V126.23.5085.5085</u>
- Phuoc, V., Sandoval-Sus, J., & Chavez, J. C. (2019). Drug therapy for double-hit lymphoma. *Drugs in context*, *8*, 2019-8-1. <u>https://doi.org/10.7573/dic.2019-8-1</u>
- Santra, A., Kumar, R., Reddy, R., Halanaik, D., Kumar, R., Bal, C. S., & Malhotra, A. (2009). FDG PET-CT in the management of primary breast lymphoma. *Clinical nuclear medicine*, 34(12), 848–853. <u>https://doi.org/10.1097/RLU.0b013e3181becdfc</u>

