

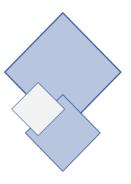


Pregnancy and Breast Cancer

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In this lecture we will go through:



1) First: 5 cases and questions about their management

2) Second: the main discussion



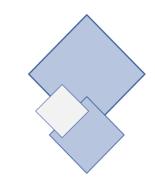
3) Third: Answers to the questions



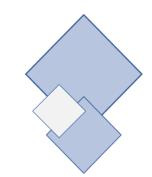
Dear audience, please type your answers in the CHAT

- 42 yr female
 - 32 weeks pregnant
- Felt a lump in right breast
- FH-, PMH-
- Exam:
 - mobile, firm, lobulated mass
 - Plan?
 - Ultrasound (US):
 - circumscribed, bilobed mass, 24 mm, BIRADS II
 - Next plan?

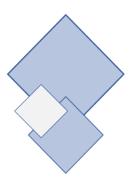




- 30 yr female, 9 w pregnant
 - Irregular mass in left breast, 2 cm
 - LAP in left axilla
 - -FH-
 - History of fibroadenoma excised at 22 years of age
 - Plan?
 - US:
 - Right breast normal, in left breast:
 - Mass with irregular border, 18 mm
 - Reactive LN in left axilla
- Next plan? Any imaging?





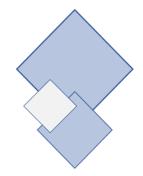


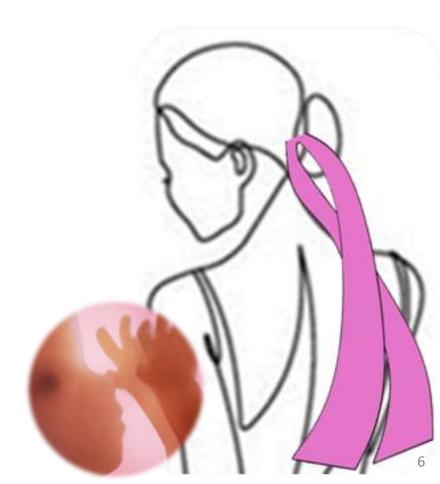
- 30 yr female, NVD 6 months ago, lactating
 - Redness and edema in UOQ and LOQ of left breast
 - No systemic signs
 - No palpable mass
 - Plan?



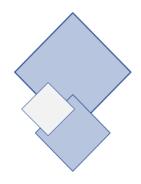
Case 4

- 25 yr female
 - Had a mass from 1 year ago in breast US
 - 7 mm, round mass
 - No previous histology
 - Now, 25 w pregnant
 - Plan?
- Follow up US:
 - 17 mm, round mass
 - Next plan?



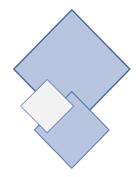


- 37 yr female, G 0
 - Underwent breast cancer treatment
 - -T1N1, ER+, PR+, HER2-
 - Did not undergo fertility preservation
 - Surgery, Chemotherapy, Radiation
 - Ended 21 months ago
 - Now under tamoxifen
 - She wants to get pregnant
 - Your recommendation?





Now, Three main topics to Discuss



1. Relation of pregnancy/lactation and breast cancer

2. Breast cancer during pregnancy

3. Pregnancy after breast cancer

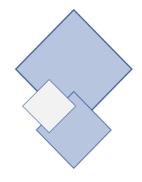
Abbreviations

breast cancer = BC

pregnancy= Py

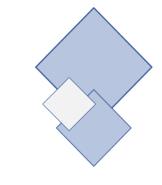
breastfeeding= BF

post-partum= PP



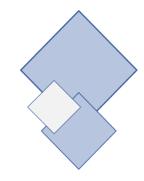
1. Relation of pregnancy and breast cancer

Py & BF: Risk or Protective Factors for BC?



- & A dilemma
 - Pregnancy and lactation: protective against BC But
 - Small increase in incidence of BC up to 2 or 5 after delivery
 - >> Cumulative effect: favorable
 - Overall, reduction in BC risk proportional to
 - the number of full-term pregnancies
 - the total duration of lactation

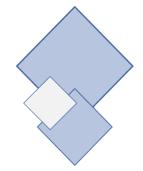
yrs





2. Breast cancer during pregnancy

Breast cancer during pregnancy Definition of PABC



PABC= Pregnancy Associated BC

Pregnancy

Breastfeeding

OR

Pregnancy 1 year postpartum

BREAST CANCER

Breast cancer during pregnancy Epidemiology of PABC





Solution Providence rate: 17 to 40 per 100,000 births

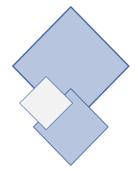
▶ But lower in Py (3.0 - 7.7) than in PP (13.8 - 32.2)

✓ PABC incidence: increasing in many countries

Maybe due to higher age of Py



Breast cancer during pregnancy Prognosis of PABC



- Delay in diagnosis
 - **▶** Later presentation
 - ✓ Higher stage at presentation
 - Treatment delays

AND

Increased proportion of receptor negative cancers

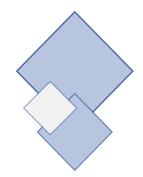
- Therefore, overall
 - Negative effect on outcome
 - Decreased overall survival



Breast cancer during pregnancy Clinical presentation of PABC

The most common

- A persistent, palpable, painless breast mass
 - Increasing in size disproportionately to the surrounding evolving breast tissue
- Skin thickening
- Asymmetric breast swelling
 - Patients *and* physicians may wrongly think these findings are normal physiologic changes!
 - Normal physiologic changes can make palpation of a mass more difficult and cause a delay in diagnosis



Breast cancer during pregnancy Diagnosis of breast lumps in Py

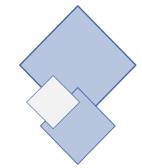
- **Uncludes**
 - → A detailed history
 - Physical exam
 - Diagnostic breast imaging: ultrasound
 - >> If suspicious for malignancy
 - ✓ Mammography ←
 - ✓ Core needle biopsy

Core needle biopsy is completely safe in Py and BF, and should be performed with any level of suspicion

In parentheses

- Benign breast lumps can enlarge during Py and BF Fibroadenoma is the most common benign solid mass in young women
 - >> Frequently enlarges during Py and BF
 - ✓ May be observed
 - ✓ Surgery rarely needed, except for very fast enlargement
 - ✓ Or very large size
 - But surely needs to be confirmed by histology

Ultrasonography



First-line imaging in Py/BF

Due to safety, and useful information

•When doctor is uncertain: US can confirm there is no lump, just NL breast tissue

•Diagnoses simple cystic lesions

•Investigates solid and atypical cystic lesions

•Gives precise description and Bi-Rads classification

Mammography

Not performed when <u>un</u>necessary

»But done if persistent doubt after US

>> often helpful and not dangerous

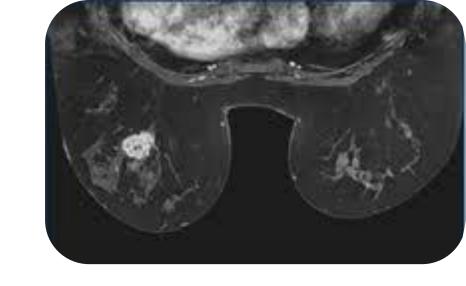
✓ If BC detected in CNB in Py/BF, bilateral mammography is necessary

With abdominal shield



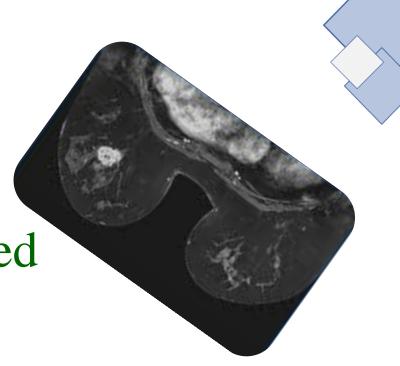
MRI in Pregnancy

- Prone position
 - >> difficult for pregnant woman
- SGadolinium enters fetal blood
 - Adverse effects seen in animals
- **%**Heating
 - can affect cell migration in 1st trimester
- **%**Noise
 - may harm fetal hearing (~24 w)
- MRI without Gadolinium may be OK
 - ≫But not helpful
 - ✓ New studies on the way

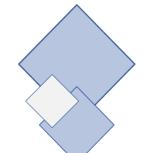


MRI in breastfeeding

- Can be performed during BF
 - Main indication: diagnosed BC
 - ▶ BC extension may be underestimated
- Some Little Gad. excreted in milk
 - ► Absorbed by infant
 - ✓ no reported cases of direct toxicity
 - 12- 24 h BF pause preferable



Breast cancer during pregnancy Metastasis work up for PABC



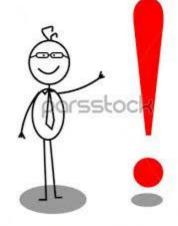
Staging in non-PABC		Staging in Py	
CT scan of abdomen	I	CXR with shield	
CT scan of chest	OR PETscar	US of liver	
bone scan	n	No bone scan	

If suspicious to bone mets: may MRI without contrast

Any suspicion for distant metastasis: needle biopsy (because metastases change treatment plans)

Breast cancer during pregnancy Diagnosis- Must Do Points:





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Physicians should have a low threshold for imaging of a persistent or enlarging breast finding in a pregnant or lactating patient

Breast cancer during pregnancy Treatment of PABC

Invasive BC



In non-metastatic BC, treatment involves:

TYPE?

✓ surgery

TIMING? // ± adjuvant or neoadjuvant chemotherapy

 \checkmark ± adjuvant endocrine therapy

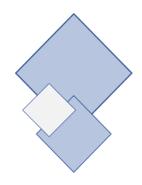
✓ ± adjuvant radiotherapy

In metastatic BC, involves:

✓ Systemic therapy < TIMING!?!

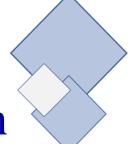


TIMING?





Breast cancer during pregnancy Treatment of PABC



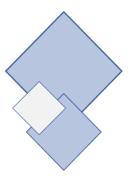
Invasive BC: treated in a multidisciplinary approach



- Termination of Py does not improve outcomes
 - should not be recommended
 - >> may safely undergo many local and systemic treatments while maintaining Py
 - ► Early delivery exceptionally considered if it affects maternal oncologic outcome



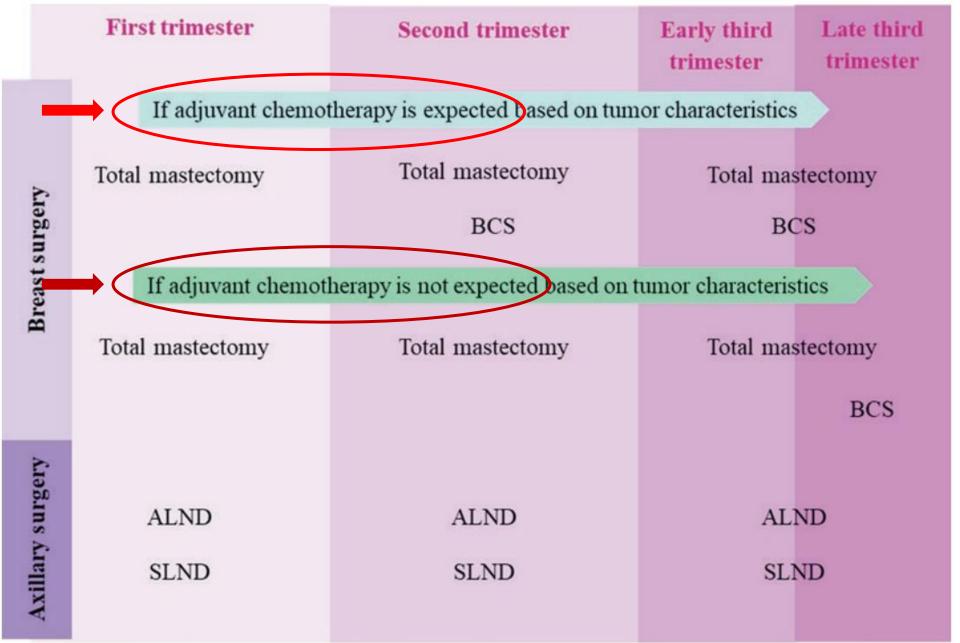
Breast cancer during pregnancy Surgery

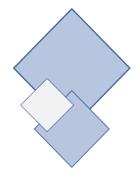


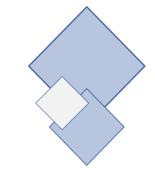
Early Py is a contraindication for breast conserving surgery (BCS)

Mastectomy is preferred in the first trimester

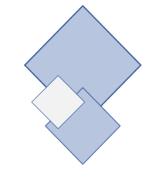
To avoid the impact of delaying radiation



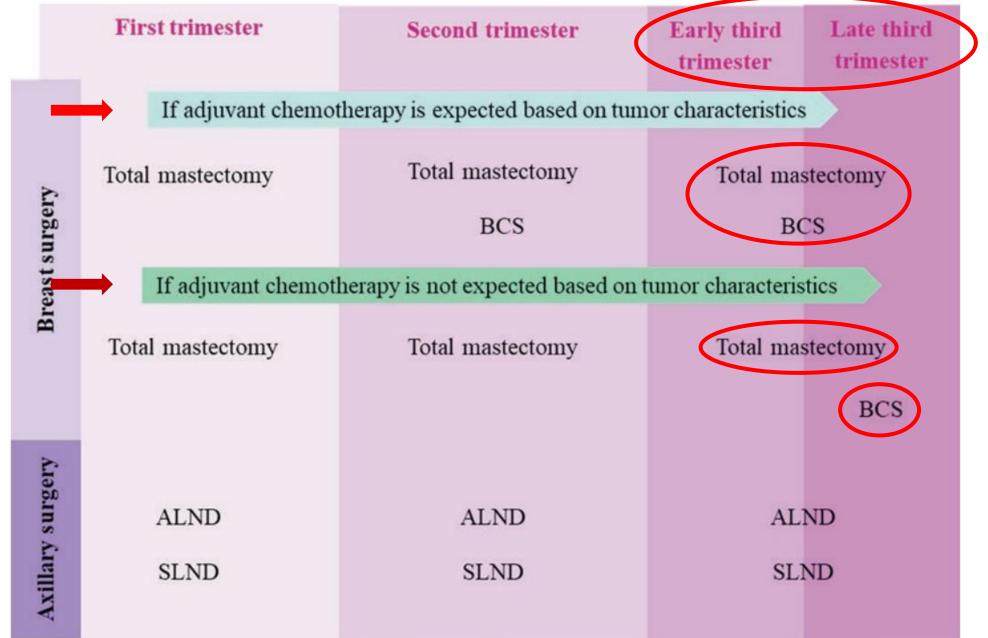


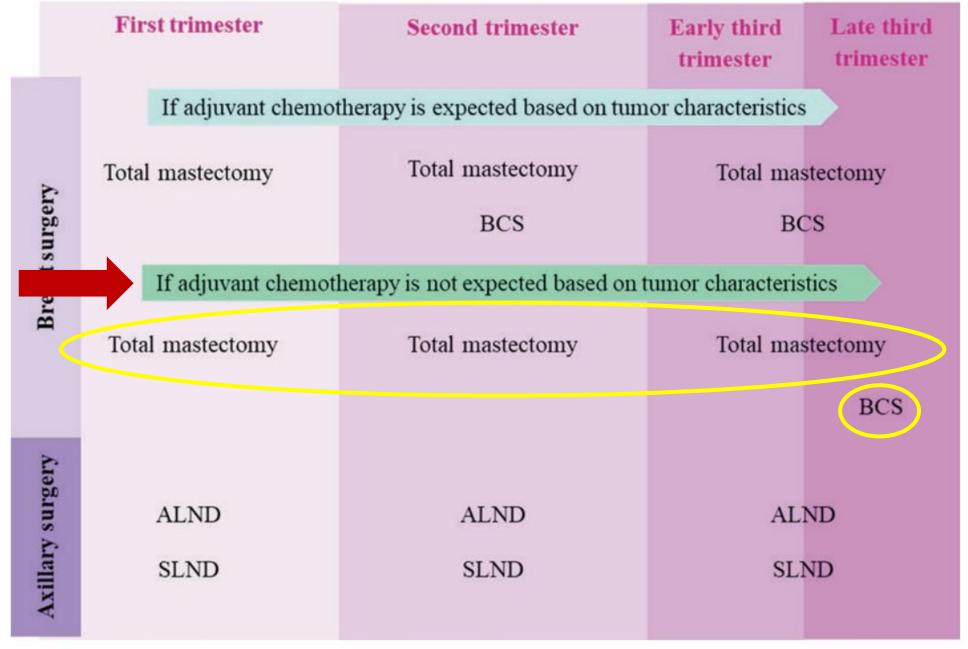


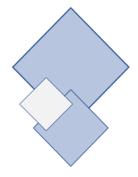
	First trimester	Second trimester	Early third trimester	Late third trimester					
•	If adjuvant chemotherapy is expected based on tumor characteristics								
Breastsurgery	Total mastectomy	Total mastectomy	Total ma	stectomy					
		BCS	ВС	CS					
east s	If adjuvant chemotherapy is not expected based on tumor characteristics								
Br	Total mastectomy	Total mastectomy	Total ma	stectomy					
				BCS					
Ţ.									
Axillary surgery	ALND	ALND	AL	ND					
illary	SLND	SLND	SL	ND					
Ax									



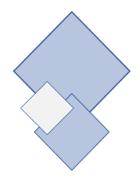
	First trimester	Second trimester	Early third trimester	Late third trimester		
	If adjuvant chemotherapy is expected based on tumor characteristics					
Breastsurgery	Total mastectomy	Total mastectomy BCS	Total mas			
east	If adjuvant chemotherapy is not expected based on tumor characteristics					
Bı	Total mastectomy	Total mastectomy	Total mas	stectomy		
				BCS		
rgery						
ıns /	ALND	ALND	AL	ND		
Axillary surgery	SLND	SLND	SL	ND		







		First trimester	Second trimester	Early third trimester	Late third trimester
		If adjuvant chemotherapy is expected based on tumor characteristics			
	Breastsurgery	Total mastectomy	Total mastectomy BCS	Total mas	
		If adjuvant chemotherapy is not expected based on tumor characteristics			
	B	Total mastectomy	Total mastectomy	Total mas	stectomy
					BCS
	Axillary surgery	ALND SLND	ALND SLND	AL:	



Breast cancer during pregnancy Axillary Surgery on PABC

In the past, only axillary dissection in PABC

Secause axillary involvement > 50%

Now increasing number of node—negative

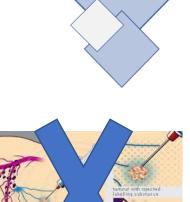
✓ might benefit from SLNB

Blue dyes are FDA category C during Py

• Radioisotopes are approved in Py

The one-day protocol is preferred

o To reduce time of isotope exposure

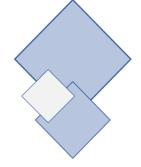






Breast cancer during pregnancy-Systemic/Adjuvant Treatments in

PABC



Main ones:

- **Solution** Chemotherapy
 - Not done in trimester 1
 - Performed in trimester 2 and 3
 - ✓ Safe interval between last ChT and elective delivery:
 - 3 weeks (or avoidance of ChT after 35w of GA)
- Sendocrine therapy, Anti-her2 therapy and Radiation
 - Not done during Py, postponed to after delivery

Breast cancer during pregnancy Inflammatory breast cancer (IBC)

- 21% to 5% of all BC

 Extremely aggressive
- Limited literature about treatment during gestation Treatment based on current guidelines
 - >> systemic chemotherapy
 - ✓ followed by mastectomy with axillary dissection
 - Radiation therapy after delivery
- There is no evidence that early termination improves the outcome.

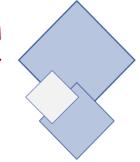
However, given the poor prognosis of IBC, patients should be counseled on the risks and benefits of continuing or terminating an early Py

In parentheses



- Mastitis and abscess are differential diagnoses for IBC
 - Much more common than IBC
 - >> US might be helpful
 - ✓ If a suspicious mass or distortion is seen
 - But US not always helpful
 - **▶** If medical treatment is not effective
 - ✓ Biopsy is needed
 - Or if there is an abscess
 - ✓ Biopsy of the wall needed at time of drainage

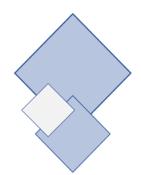
${\tt Breast\ cancer\ during\ pregnancy\ } Lactation\ during\ and\ after\ BC$



- Very limited data is available
 - Separate Beauting BC treatment including
 - Chemotherapy, endocrine therapy, targeted therapy,...
 - Signature Survivors: Signature Sign
 - **▶** BF is allowed



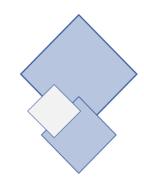
✓ Even if the milk is insufficient in the radiated breast





3. Pregnancy after breast cancer

3. Pregnancy after breast cancer

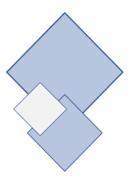


Let Is pregnancy after breast cancer treatment dangerous for the prognosis of the cancer?

Sels a concern for patients and physicians

- Limitations for prospective study
- ▶ One major bias: the "healthy mother effect"
 - ✓ survivors who are in better health or have had favorable pathologies opt for Py

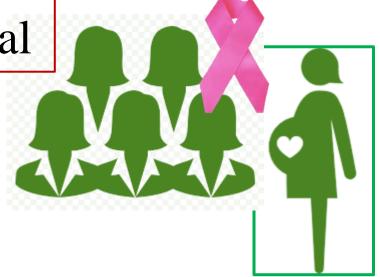
3. Pregnancy after breast cancer





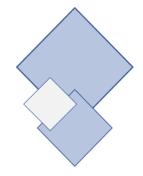
Overall survival and disease free survival





Also for ER+ tumors

Rate of obstetrical outcomes in BC survivors:



Increased incidence of

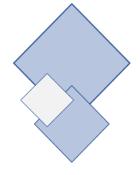
%preterm labor

Solow birth weight



sintrauterine fetal anomalies

3. Safe time interval from BC to Py



Many oncologists prefer to wait 5 years!

Breast cancer treatments

Two
years
wait

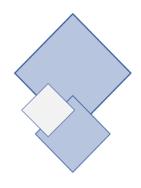
Hold
Endocrine
Therapy

Pregnancy

Continue Endocrine Therapy

Maybe because of highest recurrence in the first 2 years





Answer to cases

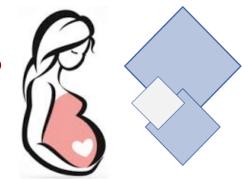


CASE 1: 42 yr, 32 w, circumscribed, bilobed mass

- -US:
 - circumscribed, bilobed mass, 24 mm, BIRADS II
 - Next plan?
 - CNB:
 - Fibroadenoma
 - Next plan?
 - Follow up with US

CASE 2: 30 yr, 9 w, 18mm irregular mass

- Next plan? Any imaging?
 - Mammography:
 - Spiculated mass in UOQ of left breast
 - Next plan?
 - CNB of mass:
 - Invasive ductal carcinoma, G2
 - ER+, PR+, Her2-
 - Next plan?
 - Surgery:
 - Mastectomy + SLN, then CHT



CASE 3, 30 yr, lactating, redness and edema

- Plan?
 - US: edema and tissue distortion in UOQ and LOQ, maybe mastitis
 - Antibiotics
 - No response
 - Plan?
 - CNB from distortion \pm skin:
 - Invasive ductal cancer, ER-, PR-, HER2-
 - Diagnosis?
 - Inflammatory breast cancer,
 - Metastasis work up: negative
 - Treatment?
 - Neoadjuvant ChT, Mastectomy, Radiation



Case 4, 25 yr, 25 w, previous 12 mm mass

- Follow up US: 17 mm, round mass
 - Next plan?
 - CNB:
 - Invasive lobular cancer, G3, ER+, PR+, Her2-
 - Next plan?
 - Surgery
 - Breast conserving surgery + SLN
 - Then ChT, then radiation after delivery

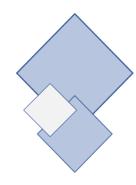


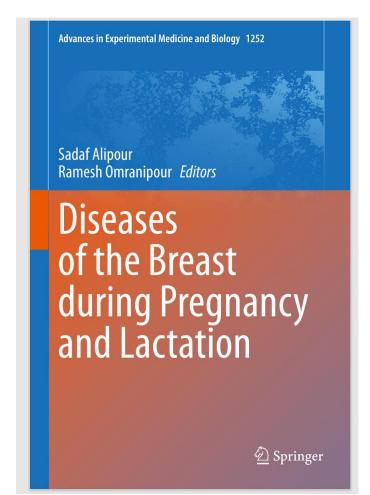
CASE 5, 33 yr, G0, BC treatment ended 21 m ago

- Now under tamoxifen
 - She wants to get pregnant
 - Your recommendation?
 - If no other problem other than the beast:
 - Hold tamoxifen temporarily after 2 years
 - Begin tamoxifen again after pregnancy

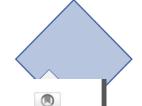
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 - Chapter 11- Clinical Presentation, Diagnosis and Prognosis of Pregnancy-Associated Breast Cancer James Sun and Marie Catherine Lee
 - Chapter 12- Surgery for Pregnancy-Associated Breast Cancer- Ramesh Omranipour
 - Chapter 15- Systemic Treatments in Pregnancy-Associated Breast Cancer-Omid S. Tehrani
 - Chapter 16- Radiotherapy in Pregnancy-Associated Breast Cancer Farnaz Amouzegar Hashemi
 - Chapter 20- nflammatory Breast Cancer in Pregnancy and Lactation Samantha Linhares, Tamrah Alrammah, Hattan A. Alghamdi, and Mecker G. Möller
 - Chapter 22- Lactation during and after Breast Cancer-Fedro A. Peccatori, Bruna Migliavacca Zucchetti, Barbara Buonomo, Giulia Bellettini, Giovanni, Codacci-Pisanelli, and Micaela Notarangelo
 - Chapter 23- Pregnancy in Breast Cancer Survivors- Vesna Bjelic-Radisic, Mohsen Esfandbod, and Sadaf Alipour
 - Chapter 27- Pregnancy and Lactation: Risk or Protective Factors for Breast Cancer? Bruna Migliavacca Zucchetti, Fedro A. Peccatori, and Giovanni Codacci-Pisanelli





Breast cancer during pregnancy





Prenatal Care during and after BC Treatment

Prevention and Treatment of Maternal Complications

- During and after Breast Cancer Surgery
- During and after Chemotherapy

or treat maternal and fetal complications of surgery and chemotherapy in pregnancy- fetal risks [3]. (see also Chaps. 3, 4). associated breast cancer.

and balance the maternal and fetal risks. In prognosis for breast cancers [1, 2]. this chapter, we discuss about how to prevent
Interdisciplinary meetings and discussions are needed to evaluate and balance the maternal and

Prevention and Treatment of Fetal Complications

- During and after Breast Cancer Surgery
- During and after Chemotherapy

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plasma volume, oxygen consumption, glomerular filtration, and coagulation state as well as decreased gastric motility, anemia, leukocytosis, and aortocaval compression. Therefore, main-

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Physical Breast Examination in

Physical Breast Examination in Pregnancy and Lactation

Abstract

Physical exam of the breast is a very important Breast examination (BE) is a skill that needs both part of breast assessment both for breast cancer screening, and when approaching breast as non-pregnancy periods. However, physical changes that occur in the breast during these times due to hormonal effects cause altera- (BI), and breast palpation (BP) [1, 2]. tions that can on one hand conceal some pathologic disorders, and may on the other hand appear as pathologic findings while first on some key points for an accurate breast examination, and then reviews some challenging controversial findings that may be noticed during breast exam in a pregnant or lactating

Breast exam · Breast palpation · Breast inspection · Breastfeeding · Pregnancy

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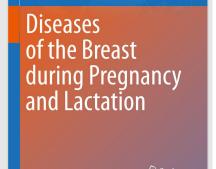
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2.1 Overview

experience and attention. One should not only put adequate time but also care enough to examlesions. Examination during pregnancy and ine all parts that are involved in breast disease, breastfeeding follows exactly the same method that is whole breasts, axillary regions, and supraand infraclavicular areas. BE includes taking related history from the patient, breast inspection

During the interview, in addition to general items such as age and queries about systemic and underlying diseases or medications, quesbeing purely physiologic. This chapter focuses tions should be asked about menstrual and reproductive history including consumption of exogenous hormones; past history of irradiation to the chest or recent trauma, any type of breast surgery, previous breast imaging, and breast symptoms including pain, swelling, skin discoloration, recent asymmetry, or enlargement; as well as nipple eczema, itching, desquamation, retraction, or secretions. Furthermore, previous history of benign breast disease and breast, ovarian or other cancer, and a family history of any of these malignancies should be enquired [3, 4].

> During BI, the examiner should inspect breasts and lymph node-bearing areas in different positions for various pathologic changes such as breast edema or asymmetry; skin edema, dimpling, ulcer, retraction, or color change; nipple



Advances in Experimental Medicine and Biology 1252

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Ramesh Ömranipour Editors



Thank you

