#### Breast Cancer



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#### Breast Cancer



- Most common cancer among women in the US
- 2nd leading cause of death in women
- 1 in 8 women will develop breast cancer
- Incidence/mortality rates have declined

#### Breast Cancer



- Breast cancer increases with age
- 95% of breast cancer cases occur in women >age 40
- Righest in white women
- 2nd highest in black women

#### Male Breast Cancer



- ≪ <1% of all breast cancer cases
  </p>
- Estrogen/androgen ratio disruption
- BRCA positive, Li-Fraumeni, Cowden's syndrome, Lynch, syndrome, Klinefelter's syndrome, hepatic dysfunction, marijuana use
- R/PR positive breast cancer

# Screening



- Clinical exam
- **Mammogram**
- R MRI
- CT scans
- R PET scans

# Staging



○ 0- DCIS○ >98% 5 year survival

I - <2 cm, negative LN</li>№ 95% 5 year survival

II- <2 cm 1-3 positive LN or 2.1 cm-5 cm if negative LN</li>≈ 85% 5 year survival

# Staging



III- Any tumor size with >4 LN, infraclavicular, ipsilateral internal mammary, supraclavicular LN, tumors >5 cm with >1 LN, chest wall or skin involvement

52% 5 year survival

≈ 18% 5 year survival

# Prognosis



- Small tumor size
- Regative lymph node
- Absence of lymphovascular invasion
- Representative Positive hormone receptor status

## Receptor Expression



- The degree of ER/PR expression is positively correlated with response to endocrine therapy
- Her2-positive status is predictive of Her2-targeted therapy



65 yo woman undergoes a routine screening mammogram which shows calcifications in the left breast that have increased since her last two mammograms



Ductal carcinoma in situ



- R Lumpectomy
- R DCIS
- R/PR positive, Her2neu positive



- Radiation and aromatase inhibitor for 5 years
- Endocrine therapy reduces the risk of developing invasive cancer in ipsilateral and contralateral breasts



- R Lumpectomy or mastectomy if disease is extensive
- Sentinel lymph node biopsy indicated if microinvasion (< 1mm of invasion) or mastectomy in whom invasion is found on final path



- Represents a heterogeneous group of neoplastic lesions confined to the breast ducts and lobules
- Oiagnosis has increased dramatically with the introduction of breast cancer screening mammography
- The risk of DCIS increases with age, is uncommon in women younger than 30 and is as high as 88 per 100,000 in women aged 50 to 64 years
- The risk of cancer-related death in women with DCIS is low, estimated at 1.9 percent within 10 years



- Most cases of DCIS are detected only on imaging studies (most commonly by the presence of mammographic microcalcifications)
- Percutaneous core biopsy under stereotactic or ultrasound guidance is preferred
- Fine needle aspiration biopsy is inadequate for the diagnosis of DCIS, as it cannot distinguish between invasive and in situ disease



- 80% of DCIS are ER/PR positive
- Consider TMX or Aromatase inhibitor (post menopausal)
- No survival benefit but decreases risk of recurrence of local breast cancer in involved and opposite breast.

#### **LCIS**



- Lobular carcinoma in situ (LCIS) is a noninvasive lesion that arises from the lobules and terminal ducts
- Represents an incidental finding that is diagnosed on a breast biopsy performed for some other reason, such as an area of fibrocystic change or a fibroadenoma

#### **LCIS**



- CR LCIS is more often detected in premenopausal than postmenopausal women, suggesting a hormonal influence in the development or maintenance of these lesions
- The cells of LCIS are typically ER-positive and have a very low proliferative rate

#### LCIS



- 4-times more likely to develop invasive cancer
- 30% risk of developing cancer over 25 years
- Republication of the Ipsilateral and contralateral breast

# Sentinel Lymph Node Biopsy



- Sentinel lymph node is the first lymph node to receive lymphatic drainage from the breast
- Lymphatic mapping done by injecting blue dye or radiolabeled tracer into the breast parenchyma near the area of cancer/mass or, alternatively, in a periareolar location
- Regional Lymph node or nodes are removed

# Sentinel Lymph Node Biopsy (DCIS)



- CR DCIS with planned mastectomy
- OCIS with suspicious features breast-conserving surgery for DCIS clinically suspected of harboring invasive cancer, including DCIS larger than 5 cm and DCIS with a palpable mass.

# Sentinel Lymph Node Biopsy (Invasive Ca)



- Early breast cancer with clinically negative node
- Fewer than three metastatic sentinel lymph nodes on SLNB
- Patients undergoing breast conserving surgery followed by whole-breast irradiation

# Axillary Lymph Node Dissection



- Three or more metastatic sentinel lymph nodes on SLNB
- One or two metastatic sentinel lymph nodes on SLNB but who do not desire whole-breast irradiation



- 32 yo female breast with a 2 cm mass in her right breast
- Mammogram confirms a 2.3 cm mass



Biopsy reveals invasive ductal carcinoma grade 3 ER/PR positive and her2 neu negative



- You refer her to a breast surgeon
- Real Lumpectomy and sentinel lymph node biopsy



- Final path reveals 2.1 cm IDC ER/PR positive, her2neu negative
- Sentinel LN is negative
- Now what?

# Gene Expression Profiles



- Oncotype Dx
- Real Endopredict
- Reast Cancer Index

## Oncotype Dx



- The Oncotype Dx 21-gene recurrence score is a prognostic assay and may identify patients who are most and least likely to derive benefit from adjuvant chemotherapy
- Indicated for women with LN negative, ER positive, HER2 negative breast cancer

## Oncotype Dx



- RT-PCR to generate a recurrence score
- Categorizes the tumor as low, intermediate or high risk
- Call Low risk- adjuvant endocrine therapy alone
  □ Tamoxifen versus Aromatase inhibitor
- ⋈ High risk- chemotherapy⋈ TC, AC-T, TAC



- Oncotype Dx recurrence score was 5, low risk
- Region TMX-5 versus 10 years
- Side effects of TMX?
- Side effects of aromatase inhibitors?
  - Retrozole, exemestane, anastrozole



- 54 yo female who presents with a very painful and swollen left breast and palpable lymph nodes
- Mammogram confirms a 6.5 cm breast mass and left axillary LN
- ™ Biopsy is c/w IDC ER/PR negative, her2neu positive
- Now what?



CT scan of chest/abd/pelvis- shows breast mass and left axillary adenopathy but no distant disease



- Neoadjuvant chemotherapy- Her2neu targeted therapy
- **Surgery**
- Radiation- Lumpectomy, axillary lymph nodes, chest wall or positive margins



- 54 yo female with history of breast cancer diagnosed 7 years ago
- Calcumpectomy, SLN followed by radiation and 5 years of Tamoxifen
- Reack pain for past 3 months
- Real Exam is unremarkable
- Treated supportively with NSAIDS, physical therapy

#### Patient Case #4



Now what?

#### Patient Case #4



- X-ray and CT scan of her spine- widespread bone mets
- Representation of the PET scan-Bone lesions but no other sites of disease
- Biopsy is done and shows IDC ER/PR positive and Her2neu negative

#### Patient Case #4



- Anastrozole plus a cyclin dependent kinase 4/6 inhibitor until progression
- Chemotherapy if any evidence of visceral crisis or progressed on multiple lines of endocrine therapy



A 50 yo woman underwent breast conservation surgery and radiation for Stage I triple negative breast cancer. She declined adjuvant chemotherapy. Two years later she is found to have a 1.5 cm ipsilateral relapse that on biopsy is again triple negative. Bone scan and CT scans are all negative



- Which of the following treatment plans would provide the greatest reduction in risk of future relapse?
- A) Mastectomy
- (R) Mastectomy with adjuvant chemotherapy
- C) Mastectomy with radiation and adjuvant chemotherapy



B) Mastectomy with adjuvant chemotherapy



A 60 yo woman with ductal carcinoma in situ was treated with breast conservation therapy, and all known disease was removed. She wants to decrease the risk of recurrent breast cancer



- Which of the following best describes the use of adjuvant endocrine for this patient?
  - A) Adjuvant endocrine therapy will decrease recurrence, regardless of whether her cancer was ER positive or negative
  - B) If her cancer is ER positive, adjuvant endocrine tx can improve OS and and decrease ipsilateral and contralateral breast recurrence
  - C) If her breast cancer is ER positive, adjuvant endocrine therapy can decrease ipsilateral and contralateral recurrence



C) If her breast cancer is ER positive, adjuvant endocrine therapy can decrease ipsilateral and contralateral recurrence



A woman presents with 3cm axillary lymph positive breast cancer, which is ER/PR negative and Her2neu negative



- Which of the following informs your decision of whether or not she should receive preoperative chemotherapy or proceed with initial surgery?
  - A) Neoadjuvant chemotherapy has been proven to improve survival in this situation
  - B) Initial surgery has been associated with improved survival in this situation
  - C) Neoadjuvant chemotherapy is associated with a reduction in the need for mastectomy and possibly axillary lymph node dissection



C) Neoadjuvant chemotherapy is associated with a reduction in the need for mastectomy and possibly axillary lymph node dissection



A 65 yo woman has a resected Her2neu positive breast cancer. She has been told that adjuvant trastuzumab/pertuzumab are beneficial



- Which of the following is the appropriate length of time for her to receive adjuvant Her2neu directed therapy?
  - A) 6 months
  - B) 1 year
  - ∝ C) 2 years
  - D) Adjuvant Her2neu directed therapy is only necessary during the time that she is receiving adjuvant chemotherapy



○ B) 1 year



- Which of the following patients would you refer for genetic counselling?
  - a) Men with breast cancer
  - b) Women diagnosed at or under the age of 50, or with Ashkenazi heritage, and/or a strong family history of breast or ovarian cancer.
  - c) Women with triple negative breast cancer up to age 60
  - d) All the above



○ D) All the above