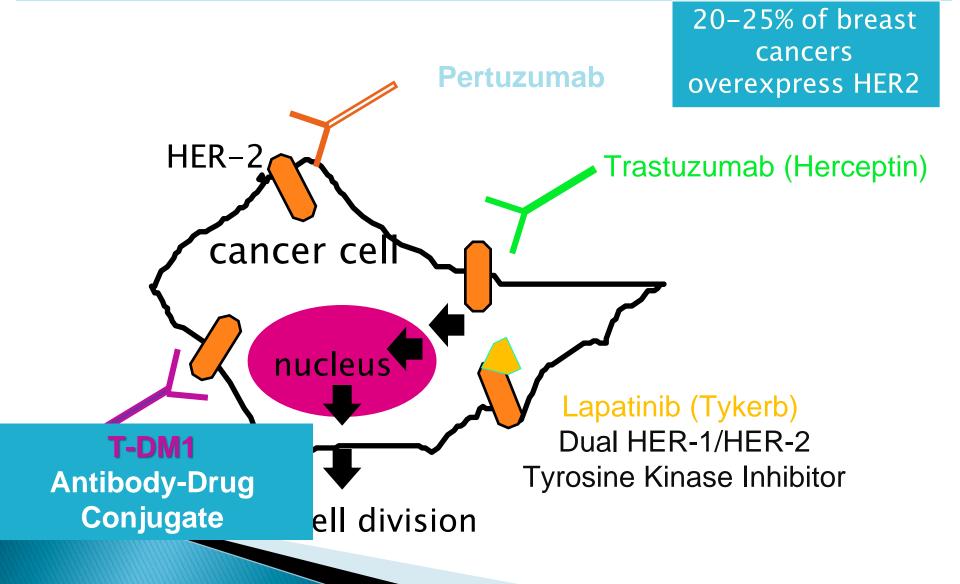


Monoclonal antibodies targeting a HER2 protein overexpressing cell

Four US FDA-Approved Drugs with HER-2 as a Target



Adjuvant (Early Stage) HER-2 Targeted Therapy

- Anti-HER2 monoclonal antibody trastuzumab (Herceptin) for 1 year is standard
 - Reduces recurrence by 1/2 & deaths by 1/3 when added to chemo in early stage breast cancer
 - Trastuzumab going off patent soon, and prices will drop
- All regimens include chemotherapy in addition to HER2 targeting therapy

European School of Oncology Guideline: HER2 Targeted Therapy for Metastatic Breast Cancer

Cardosa F et al, J Natl Cancer Inst 101:1174-1181, 2009

- Anti-HER2 therapy should be offered <u>early</u> to all HER2+ metastatic breast cancer patients unless contraindicated (or unavailable)
- Optimal duration of anti-HER2 therapy for metastatic breast cancer (when to stop) unknown

Trastuzumab/Herceptin

Given to patients whose cancer cells overexpress Her-2-neu as measured by IHC or FISH (25 to 30% of patients)

Contraindications for Breast Conserving Therapy

- Absolute:
- Prior radiation to the breast or chest wall
- Pregnancy
- Muticentric disease
- Diffuse, malignant appearing microcalcifications

Relative Contraindications for BCT

- History of collagen vascular disease
- Very large tumor > 5cms
- Very large breasts

Incremental Benefit of Adjuvant Treatments in Early Stage Breast Cancer in USA

chemotherapy + endocrine therapy + HER2 targeted therapy

chemotherapy + endocrine therapy

mastectomy

No surgery

Survival

Carcinoma in pregnancy

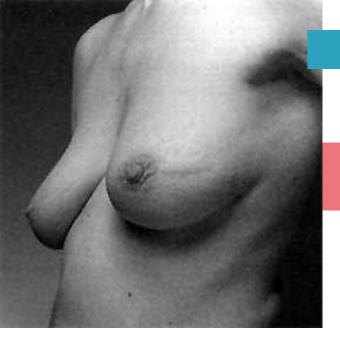
THEY ARE DIAGNOSED LATE
THEY BEHAVE THE SAME AS NON-PREGN.
THEY ARE TREATED THE SAME
NO BREAST CONSERVING SURGERY
NO RADIOTHERAPY; TERATOGENIC
NO CHEMOTHERAPY IN THE FIRST TRIMESTER
NO HORMONAL THERAPY; BECAUSE
THEY ARE HORMONE RECEPT.-VE

BRCA1/BRCA2?

- No "treatment"
- Mastectomy cuts risk by 90%
- Hysterectomy slashes another 50%
- Tamoxifen
 - anti-estrogen drug
 - Lowers risk of diagnosis of benign tumors Influenced by estrogen
 - Reduces risk about 28%
 - May reduce need for biopsies in high-risk women
 - Intervenes before invasive cancer begins

Surgery

- Lumpectomy
 - removal of the cancerous tissue and a surrounding area of normal tissue
- Simple mastectomy
 - removes the entire breast but no other structures
- Modified radical mastectomy
 - removes the breast and the underarm lymph nodes



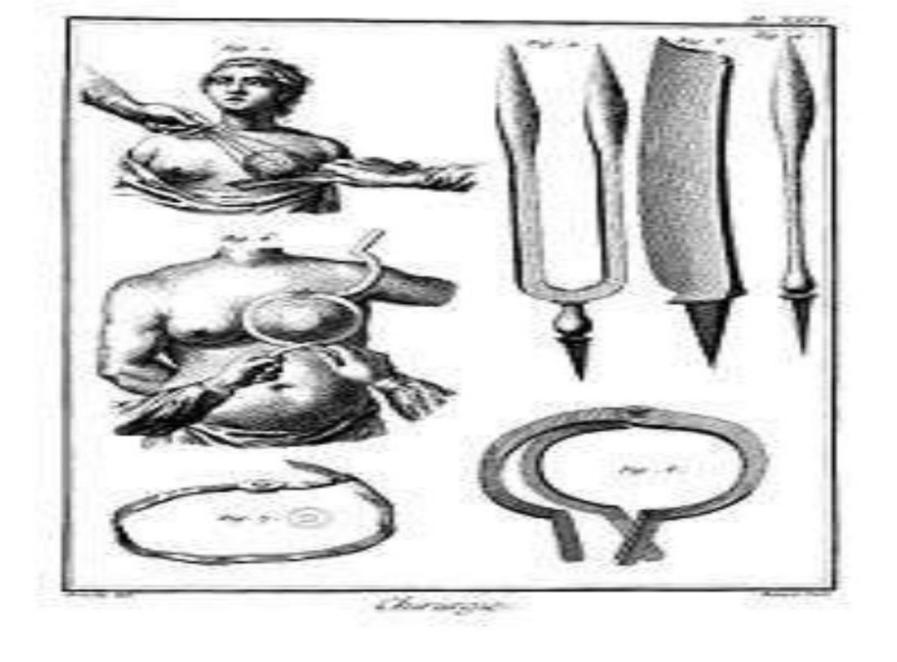
← Lumpectomy

Simple Mastectomy





← Modified Radical



Breast cancer surgery in 18th century



Breasts after double mastectomy followed by nipple-sparing reconstruction with implants

Surgical treatment of early breast cancer:

Modified radical mastectomy Pate's operation

Indications:

1st &2nd Stage of breast cancer

Retract pectoralis major at lateral border to expose pectoralis minor

