

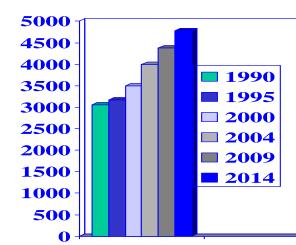




**Breast Cancer in Scotland: Current** issues/future visions Alastair M Thompson **Professor of Surgical Oncology** University of Dundee a.m.thompson @ dundee.ac.uk

#### **Breast Cancer**

- Increasingly common
- 205 to 481 new cancers in Tayside and NE Fife 2009
- ~60% to >90% 5 year survival



• Multidisciplinary evidence based practice.



### Imaging

- Breast Screening who, when, how
- Detect Cancer Early
- DCIS which will progress?
- Screening prevents 1300 deaths pa qv 4,000 unnecessary treatment (1% of 307,000 pa)
- 10,00 women, 681 cancers in 20 years, 129 overdiagnosed ,43 deaths prevented
- 20% reduction in relative risk of dying from breast cancer among invitees

### **Pathology**

- Increasing quality and quantity of information on breast cancers
- Information pre-operatively (type, ER, R, HER2)
- Attention to margins benefits for treatment planning and outcome
- Tissue banking to improve research



- Molecular pathology tumour categories, relevant to outcome and treatment response
- Pressure on services



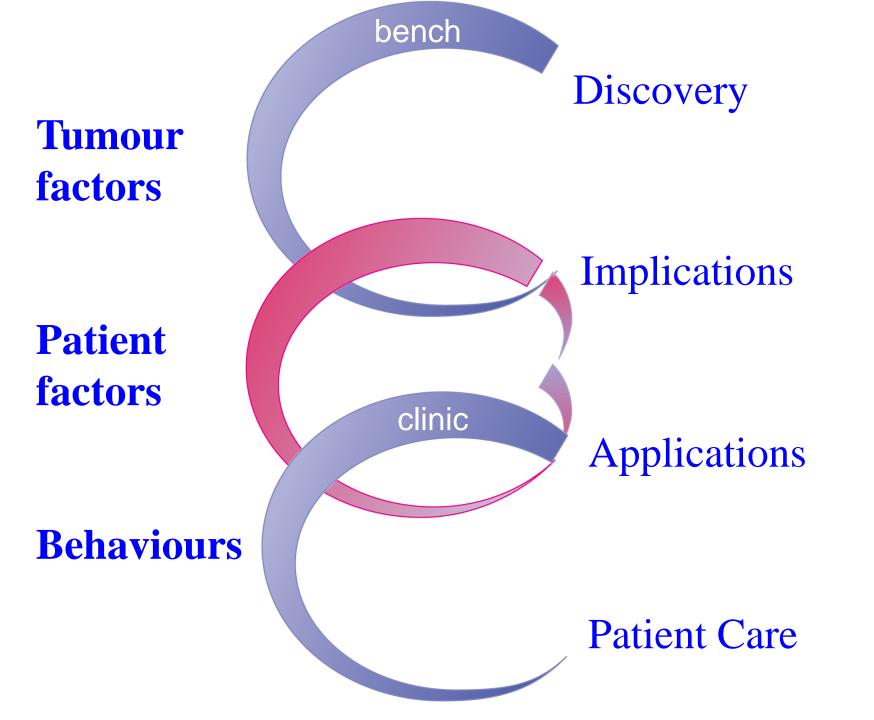
- "Less is as good as more extensive surgery"
- Mastectomy = breast conservation for survival
- Sentinel node biopsy for more patients (qv axillary node clearance)
- More exact excisions (margins)
- Oncoplastic approaches
- Better reconstruction (autologous (abdominal) flap; synthetic collagen; implants)
- Minimally invasive surgery.....

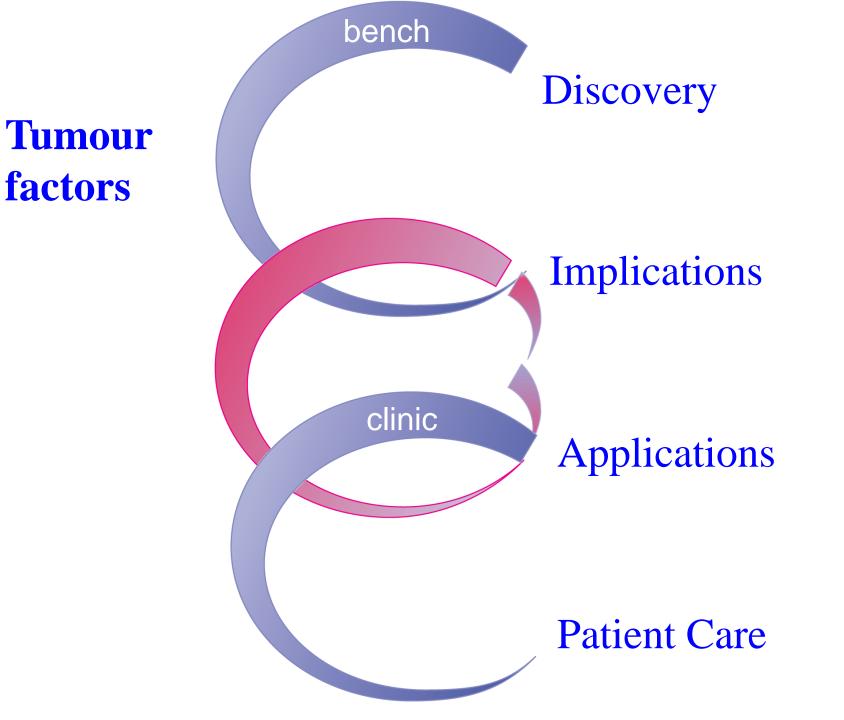
## **Drug therapy**

- Increased survival with anthracyclines, ?taxanes
- Targeted therapies HER2 (Trastuzumab, T-DM1, Pertusumab)
- Increased use of pre-operative therapy/ trial designs
- Selection of those who do and those who do not benefit from chemotherapy (MINDACT)
- Toxicities (esp of combination therapies)
- Costs v (small) incremental gain

### Radiotherapy

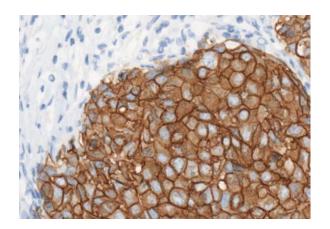
- Shorter fractionation (START Trial)
- Increasing fractionation (FAST, FAST-forward)
- Intraoperative radiotherapy (TARGIT Trial)
- Intensity Modulated Radiotherapy (IMPORT Trials)
- Conservatism of some practitioners (£)
- Inadequate resources (kit, physicists, research radiographers)

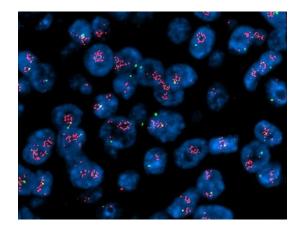




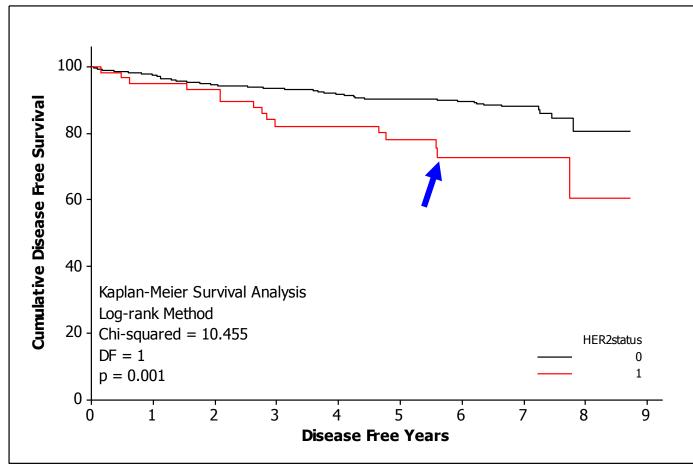
### **Undertreating ER+ HER2+ cancers**

- Capture all demographic, clinical, pathological, ER & HER2 data
- CB11 immunohistochemistry
- HER2 FISH on all 2+ cases
- n=776, population based audit

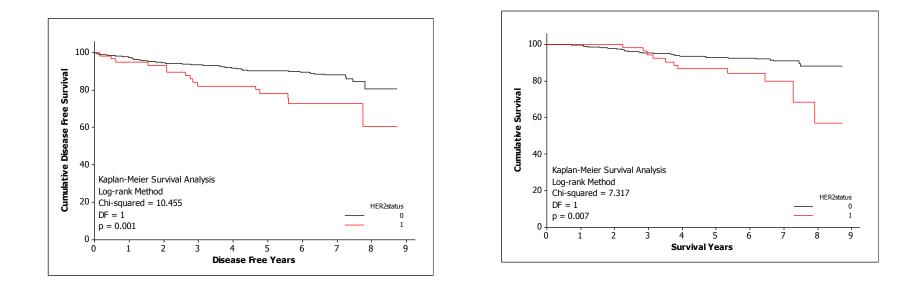




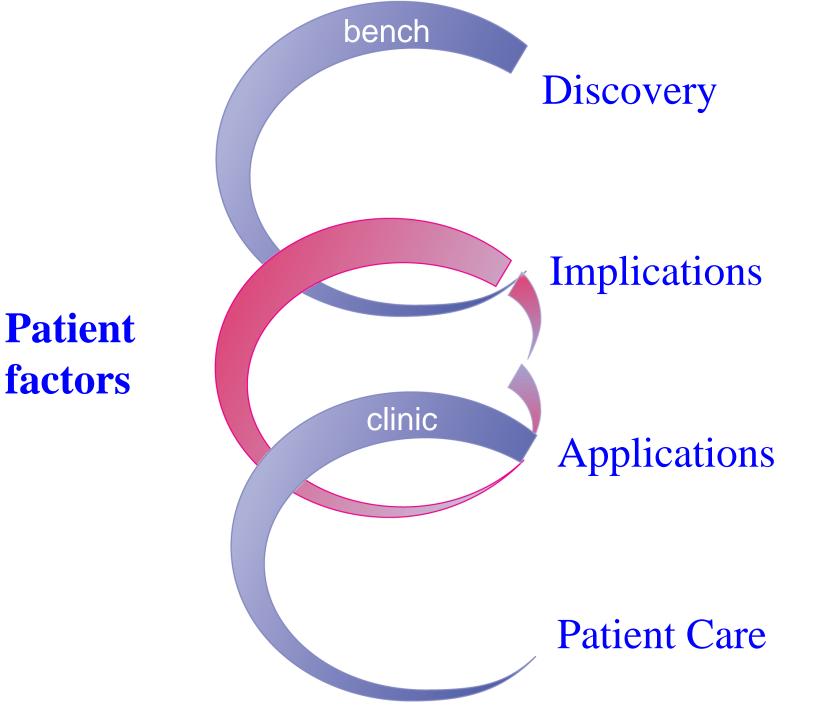
#### **ER+ HER2+ status and DFS** n=737



#### **ER+ HER2+ status and DFS/OS** n=737

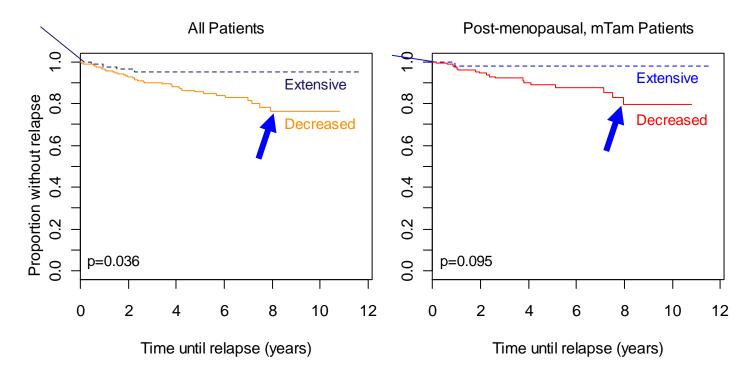


Increased mortality in HER2 positive, ER positive invasive breast cancer: a population based study. Purdie CA et al, British Journal of Cancer (2010) 103:475-81.



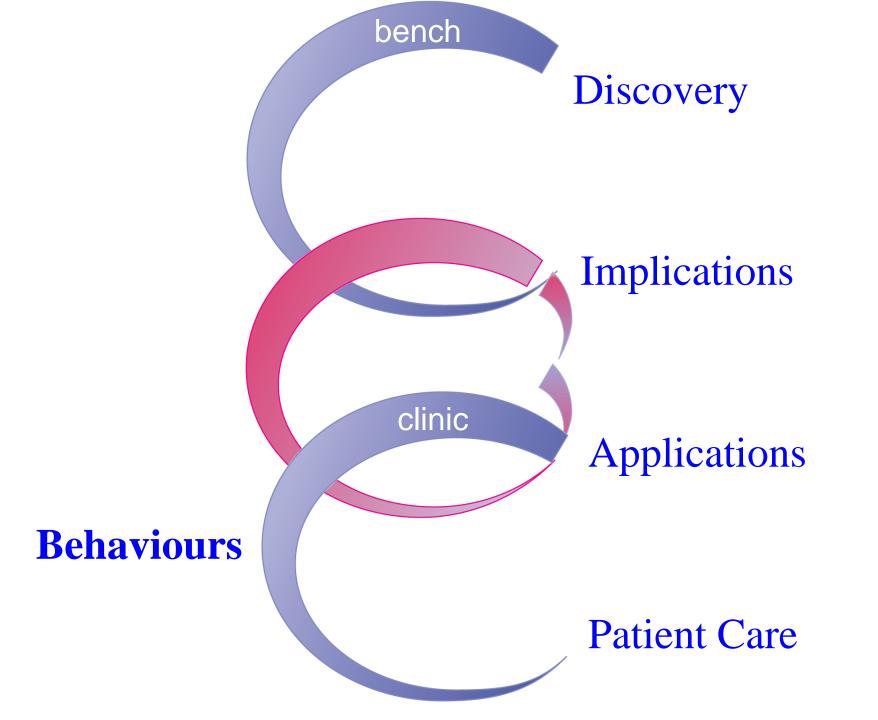
### Who should have tamoxifen?

- CYP450 2D6: tamoxifen/endoxifen
- Dundee/Manchester/Roche Diagnostics Amplichip (comprehensive analysis)



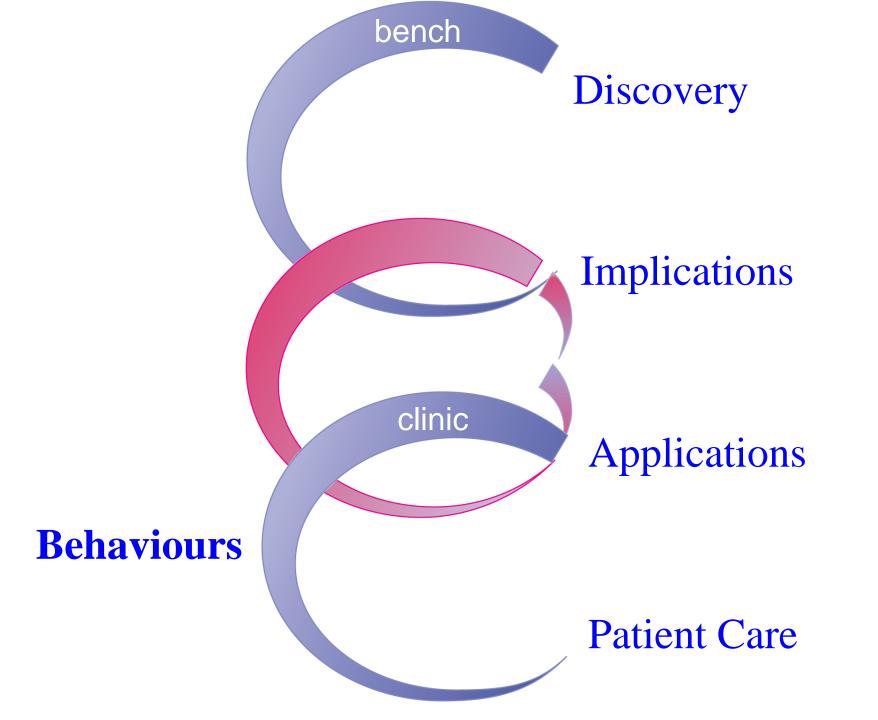
#### **CYP450 2D6**

- Comprehensive CYP2D6 genotype and adherence affect outcome in breast cancer patients treated with tamoxifen monotherapy. Thompson AM, et al. Breast Cancer Research and Treatment (2010) <u>125</u>: 279-287.
- Avoidance of CYP2D6 inhibitors in patients receiving tamoxifen. Ferraldeschi R, Howell SJ, Thompson AM, Newman WG. J Clin Oncol. 2010; 28:e584-5
- 6,000+ patient meta analysis.....



### What about adherence?

- Comprehensive CYP2D6 genotype and adherence: reassigning patients with adherence <80% to the decreased metabolizer group resulted in the HR of this group increasing.
- Tamoxifen, aromatase inhibitor adherence: adherence <80% increased mortality; younger women particularly

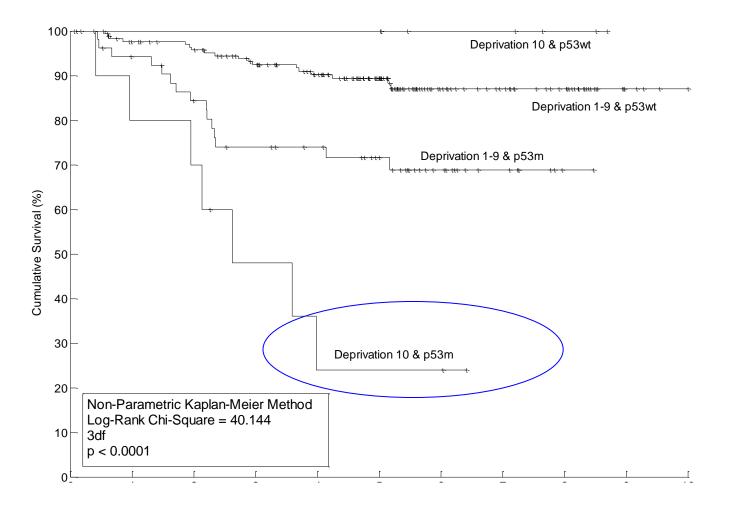


## Socioeconomic status and breast cancer

Tayside Tissue Bank/Roche p53 Amplichip Patients with *p53* mutation in the 10th decile had:

- worse disease-free survival (20% at 5 years)
- worse overall survival (24% at 5 years)
- p53 mutation, deprivation and poor prognosis in primary breast cancer. Baker L et al British Journal of Cancer (2010); <u>102</u>: 719-726.

#### **Deprivation**



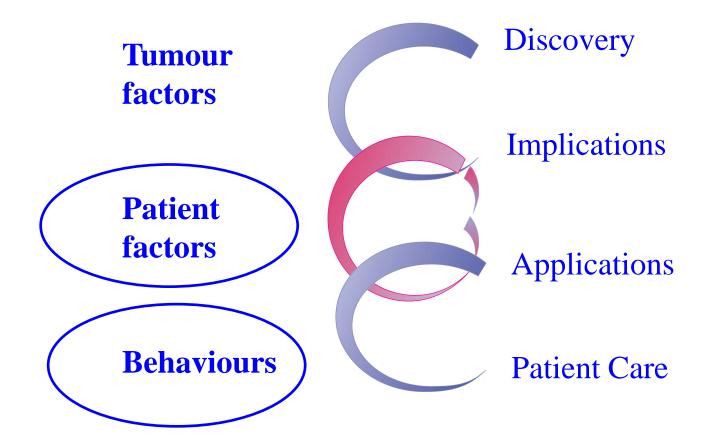


### Vision for 2013 and beyond...

An innovative scientific and clinical approach to breast cancer across Scotland (and linking beyond) advancing the:

- prevention
- early detection
- treatments and
- cure of breast cancer.....

#### What now?



## **Opportunities:**

- Primary prevention:
- Exercise, diet, alcohol, smoking
- Drugs (?)
- DCE, breast screening, workplace, GP
- Secondary (after treatment) prevention:
- Drugs (tamoxifen, aromatase inhibitors, metformin, aspirin, etc etc)
- Exercise, diet

# **Breast Cancer Research in the University of Dundee is supported**



researching the cure



DUNDEE











Almac, Aridhia, AstraZeneca, Cyclacel, Roche, Wyeth









#### a.m.thompson@dundee.ac.uk