



To improve the care of older patients with cancer around the world



# Elderly Breast Cancer – Systemic Therapy

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**Scientific Committee Meeting**

**International Society of Geriatric Oncology (SIOG 2017)**



**20<sup>th</sup> April, 2017**





*To improve the care of older patients with cancer around the world*



## Disclosures

Consultant or Advisory Role:

AstraZeneca, Aptus, Astellas, Eisai, GlaxoSmithKline,  
Foundation Medicine, Novartis & Pfizer

**Due to time constraint, I will just be focusing on systemic therapy for EBC patients!**





## Outline

- The undeniable emerging needs
- Current Dilemmas for Elderly BC patients
- Opportunities & Challenges
- Conclusion





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# Age is a known risk factor...

- **AGE** is one of the strongest risk factors for cancer development
- The aging population presents a major epidemiological challenge
  - Demographic shift brought by the baby-boomer generation



14<sup>th</sup> International Society of Geriatric Oncology, Lisbon, October, 2014  
Wedding U. Future Oncol, 2015 Mar;11(6):893-5. doi: 10.2217/fon.15.4.

# Aging Society & Aging Cancer Patients

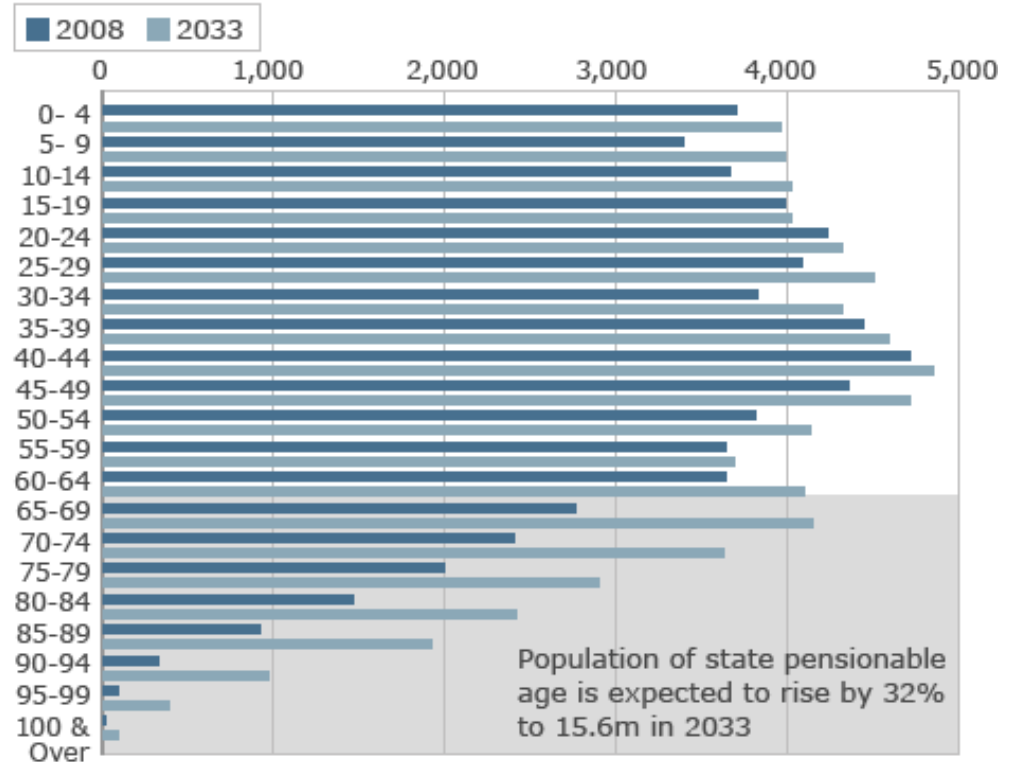
By 2030, there could be 50% more people greater than 65 years old, and 100% more people greater than 80 years old

By the year 2030, most patients with cancer will be aged over 65 years and many will be frail.



## Projected population

Expected increases in UK population by age group 2008 - 2033 (thousands)

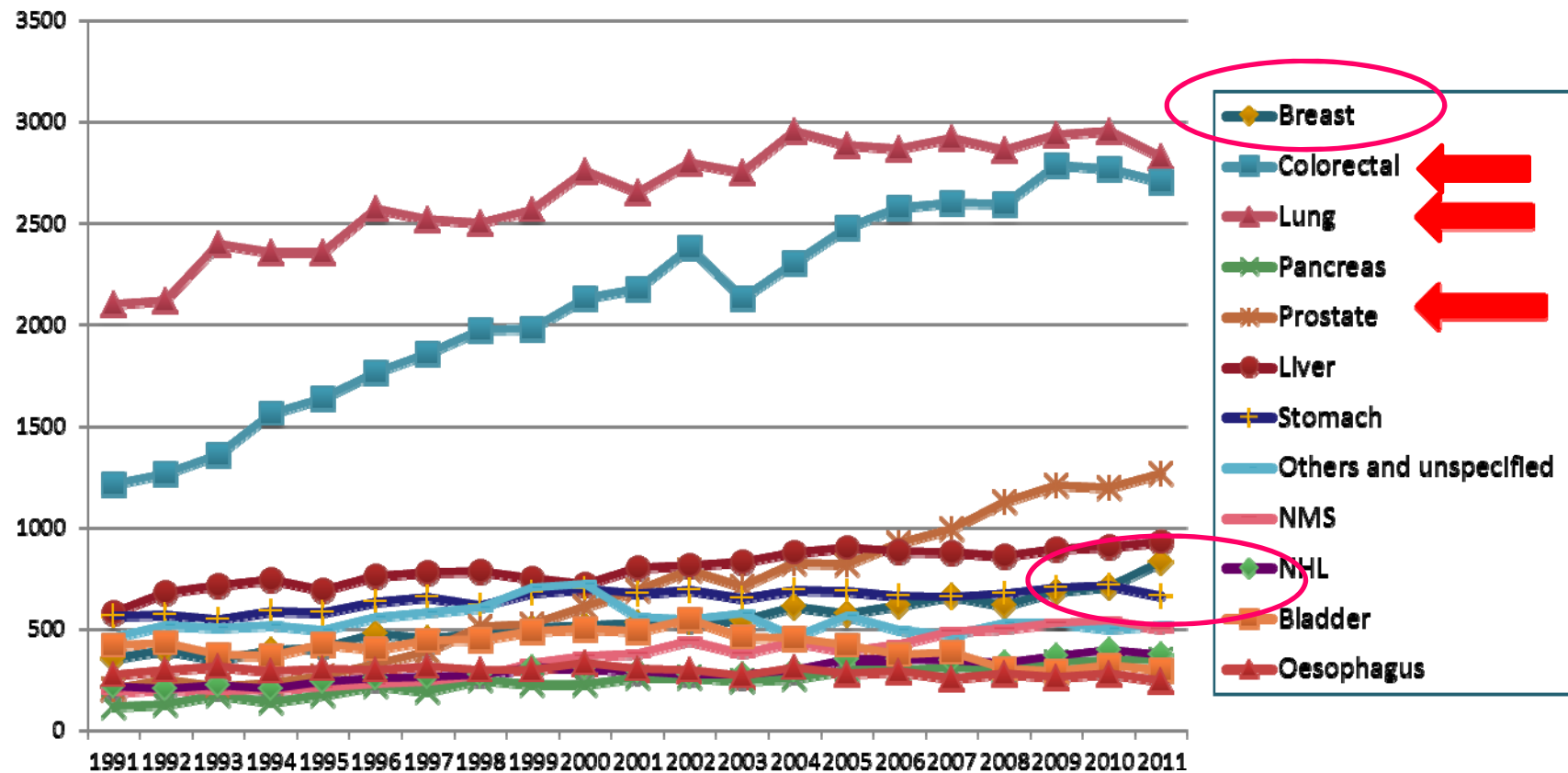


Population of state pensionable age is expected to rise by 32% to 15.6m in 2033

Source: ONS

# Age and Cancer Trends: A 20-year review of the Hong Kong Cancer Registry

## Cancer Incidence for 65 years old or above (1991-2011)



# Distribution of Female Breast Cancer by Age Group & Type in 2011

Distribution of Female Breast Cancer by Age Group and Type (Invasive and Ca in-situ) in 2011  
 2011 年按年齡組別及癌腫分類 [原位(Ca in-situ)及入侵性(Invasive)] 乳癌的發病數字

Age (yrs) 年齡(歲)	0-	5-	10-	15-	20-	25-	30-	35-	40-	45-	50-	55-	60-	65-	70-	75-	80-	85+	All ages 所有年齡
Invasive 入侵性	0	0	0	0	3	22	68	173	360	620	524	434	397	225	170	154	123	146	3,419
Ca in-situ 原位癌	0	0	0	1	0	1	9	26	66	103	84	69	42	30	22	16	7	8	484
Total 總數	0	0	0	1	3	23	77	199	426	723	608	503	439	255	192	170	130	154	3,903

年齡超過65歲之乳腺癌患者:  
**Invasive BC aged >=65":**  
**818/3419 = 23.9%**

1 in 4  
 65 y.o. or  
 above

Age-specific Incidence rates of Female Breast Cancer (per 100,000 women) by Type (Invasive and Ca in-situ) in 2011  
 2011 年按癌腫分類 [原位(Ca in-situ)及入侵性(Invasive)] 乳癌的各年齡組別發病率 (每十萬名女性人口計算)

Age (yrs) 年齡(歲)	0-	5-	10-	15-	20-	25-	30-	35-	40-	45-	50-	55-	60-	65-	70-	75-	80-	85+	Lifetime risk* 一生累積風險*	Crude rate 粗發病率	ASR (World)* 年齡標準化率 (世界)**	
Invasive 入侵性	0.0	0.0	0.0	0.0	1.3	7.2	21.0	52.7	109.1	174.0	161.3	146.8	146.8	146.8	146.8	146.8	146.8	146.8	1 in 17	90.7	61.0	
Ca in-situ 原位癌	0.0	0.0	0.0	0.5	0.0	0.3	2.8	7.9	20.0	28.9	25.9	25.9	25.9	25.9	25.9	25.9	25.9	25.9	25.9	1 in 112	12.8	8.8
Total 總數	0.0	0.0	0.0	0.5	1.3	7.6	23.8	60.7	129.1	202.9	187.2	172.7	172.7	172.7	172.7	172.7	172.7	172.7	172.7	1 in 15	103.6	69.9

1 in 8  
 75 y.o. or  
 above

年齡超過75歲之乳腺癌患者:  
**Invasive BC aged >= 75:**  
**423/3419 = 12.37%**

\* Cumulative lifetime risk before the age of 75. 一生累積風險(0-74 歲)

\*\* Rates are standardized to the age distribution of the "WHO 2000" World Standard Population. Comparisons with other countries are only valid if they use the same standard population for calculations.  
 年齡標準化發病/死亡率是根據世界衛生組織2000年標準人口計算。用於進行比較不同人群之間的疾病率時，需以同一標準人口為基礎。

Source: Hong Kong Cancer Registry, Hospital Authority  
 資料來源: 醫院管理局香港癌症資料統計中心

Nov 2013





# Elderly cancer patients are different...

- Age-related reduced organ functions
- Multiple co-morbidities
- Changes in cognition
  - Dementia, delirium
  - 1% in 65-69 y.o., 41% in >90 y.o. having dementia
- Falls
- **Poly-pharmacy**
- Higher prevalence of **depression & anxiety**

# Elderly cancer patients are different...

- Different cancer types
  - Marked increase in epithelial carcinomas from 40 to 80 y.o.
  - Cancer & aging share common etiologies – genomic instability & reduced rate of autophagy
  - **Breast Cancer among elderly are mostly hormone positive with slow tempo of disease.**

# Impact of Aging on treatment decision making...

- Surgery – definitive vs palliative
  - Surgical and anaesthetic risk
- Radiotherapy – tolerance and toxicities
  - Poor nutrition or malnourished, cachexia
- Chemotherapy
  - Dose adjustment, dose reduction, tailored regimen
- Hormonal therapy
- Targeted therapy



## Outline

- The undeniable emerging needs
- **Current Dilemmas for Elderly BC patients**
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# Misconception...

## about Elderly Cancer Patients

- “Chronological age used to be the sole important determining factor driving the aging process”
- “Chronological age is the common reference to decide treatment plan for elderly cancer patients”
- “Elderly cancer patients do not tolerate chemotherapy or any anticancer agent easily”
- “There is much toxicities related to elderly with multiple comorbidities...”

# Current Dilemmas

## 1) Therapeutic **nihilism**

- Elderly patients do not receive any treatment

## 2) The **intermediate** position?

• –Elderly patients may benefit from treatments

## 3) Blind therapeutic **enthusiasm**

- Elderly patients receive

futile/non- beneficial treatments



# Current Dilemmas

1) Therapeutic **nihilism**

- Elderly patients do

2) The **inter**

• -Eld

3)

**Under-treatment vs Over-treatment...**

...or treatments

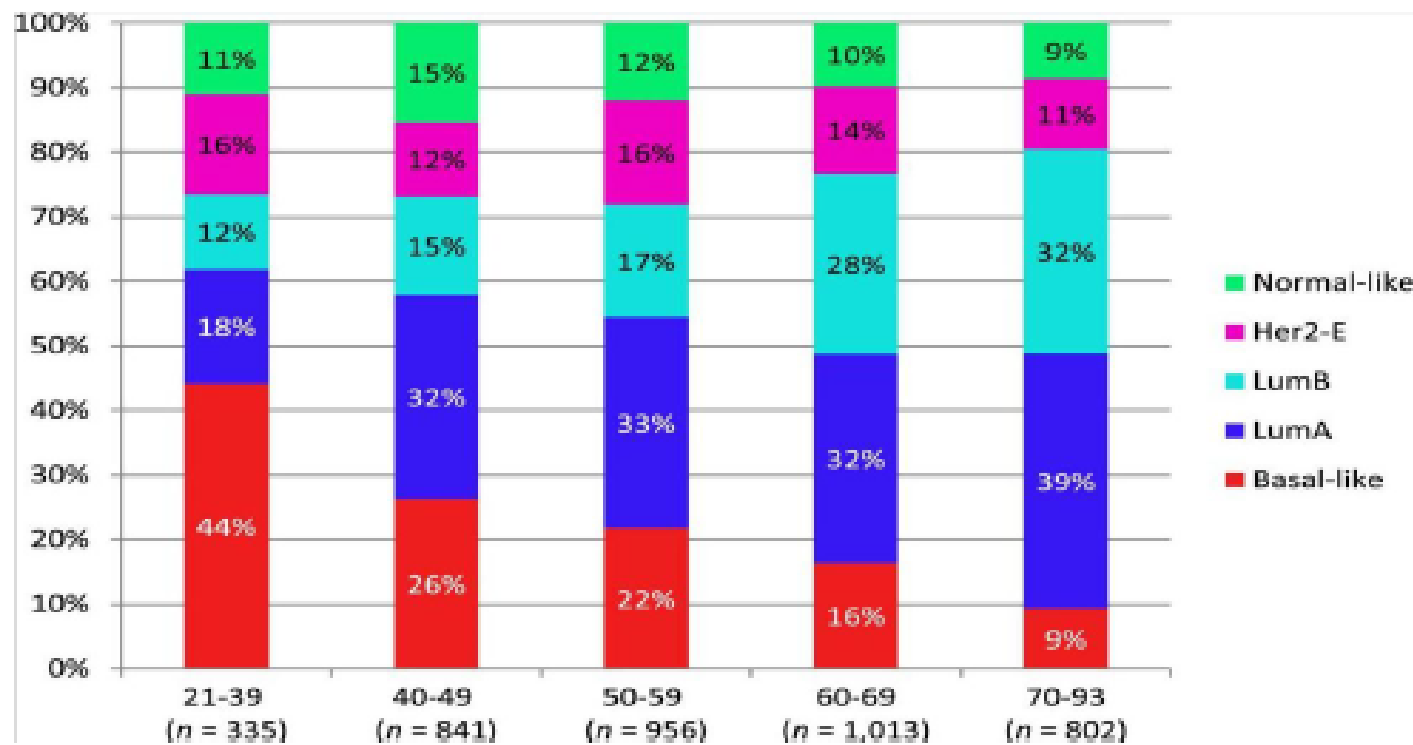


# Biology of Aging

- Almost all age-related changes lead to reduced organ function.
- There are **3 different trajectories** of aging:
  - Aging with pathology & disability
  - Normal aging with some disability
  - Successful aging with minimal disability
- **Aging is a heterogeneous process...**



# BC biology according to age

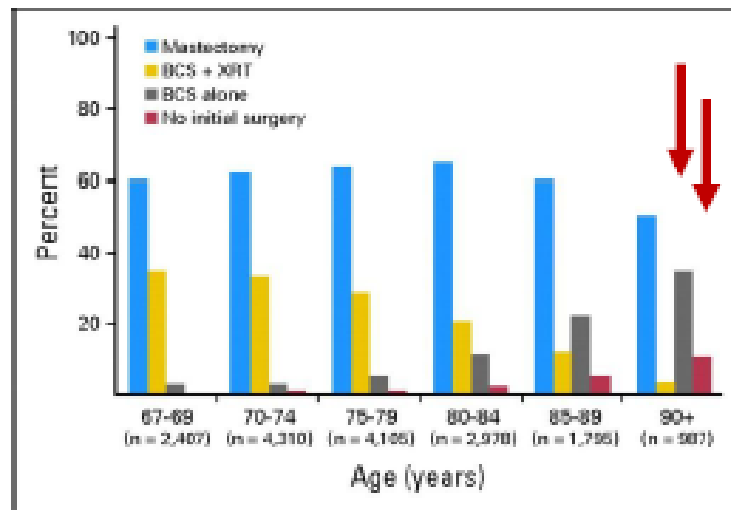


de Kruif Mol Oncol 2014, Jenskins Oncologist 2014

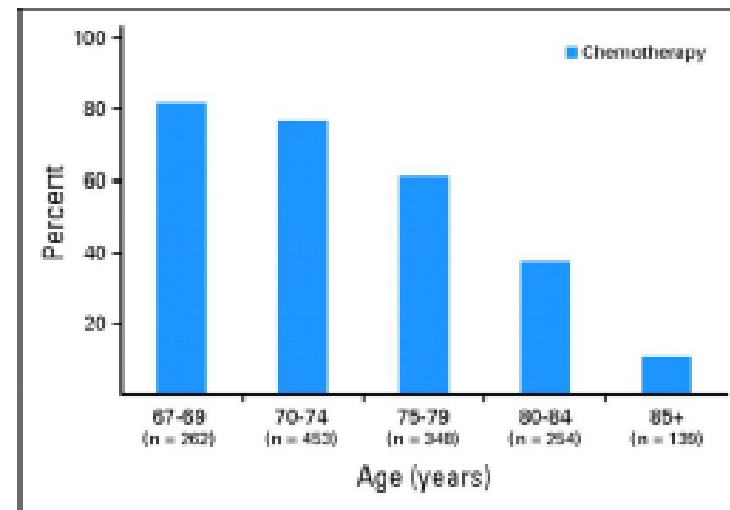
# Undertreatment

SEER database ; 49616 women with stage I/II breast cancer  $\geq 67y$

Initial treatment for stage II breast cancer by age



Treated with chemotherapy if ER+ N+ stage I/II breast cancer



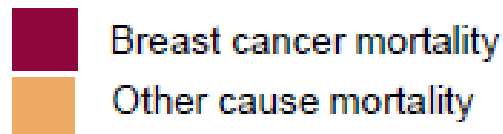
BCS = breast conserving surgery ; XRT = radiotherapy

Schonberg JCO 2010

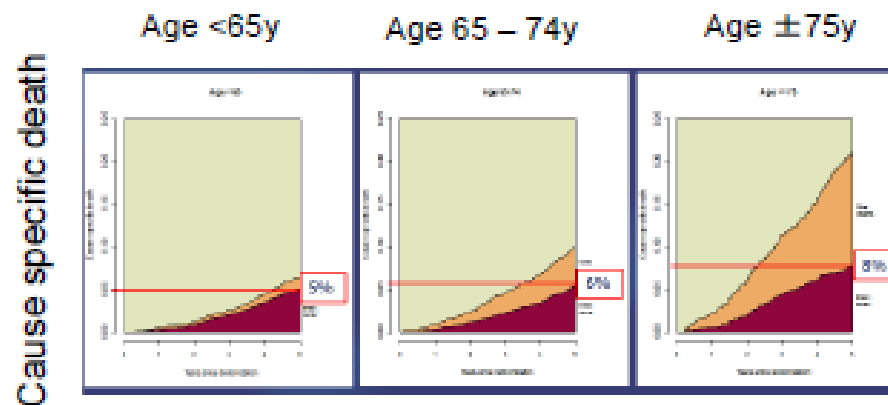
# Undertreatment

Characteristic	HR of Death Due to Breast Cancer	Range	HR of Death Due to Other Causes	Range
Age group, years	Reference		Reference	
67-69	1.0	0.9-1.2	1.3	1.2-1.4
70-74	1.1	0.99-1.3	1.9	1.8-2.2
75-79	1.2	1.1-1.4	3.0	2.7-3.3
80-84	1.5	1.3-1.7	4.1	3.7-4.6
≥ 90	1.8	1.5-2.2	5.9	5.2-6.7

Substudy from TEAM trial (adjuvant exemestane)



- Univariate HR 1.66  
(95% CI 1.34-2.08), p<0.001
- Multivariable HR 1.63  
(95% CI 1.23-2.16), p<0.001



Schonberg JCO 2010, Van de Water JAMA 2012

# Overtreatment

- A sizeable proportion of elderly with operable breast cancer die of **NON-CANCER-related causes**

N = 14048 new early breast cancer, ≥50y, FUP 4,7y

	Total deaths	Deaths from breast cancer	%
50–69	1334	933	70
70–74	514	293	57
75–79	696	329	47
≥80	1681	663	39
Total	4225	2218	53

53% of elderly BC patients aged 75-79 and 61% elderly patients aged ≥ 80 died of other non-BC causes

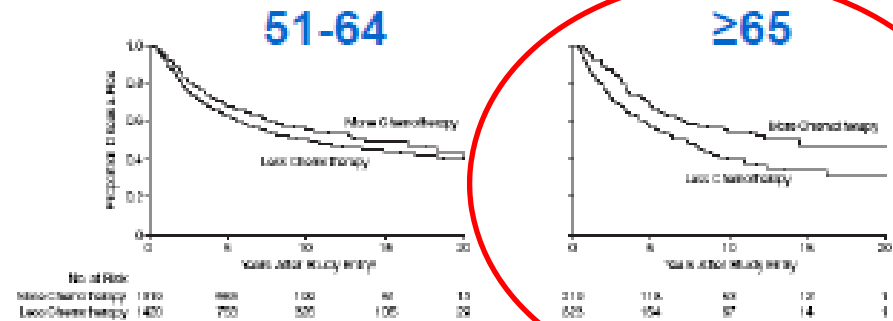
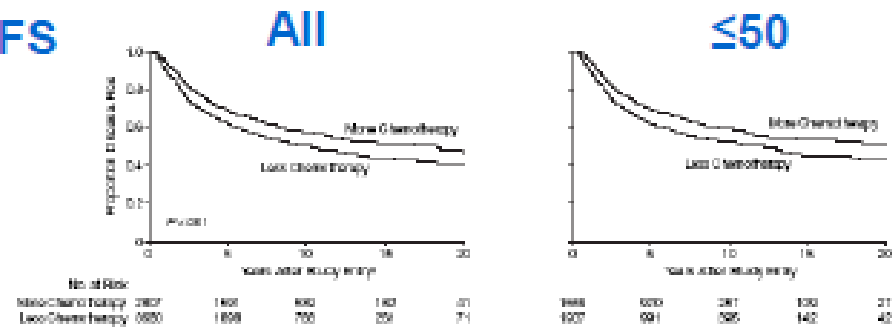


- Absolute **benefit** of treatments is **lower**

Ali Br J Cancer 2011

# Adjuvant chemo

DFS



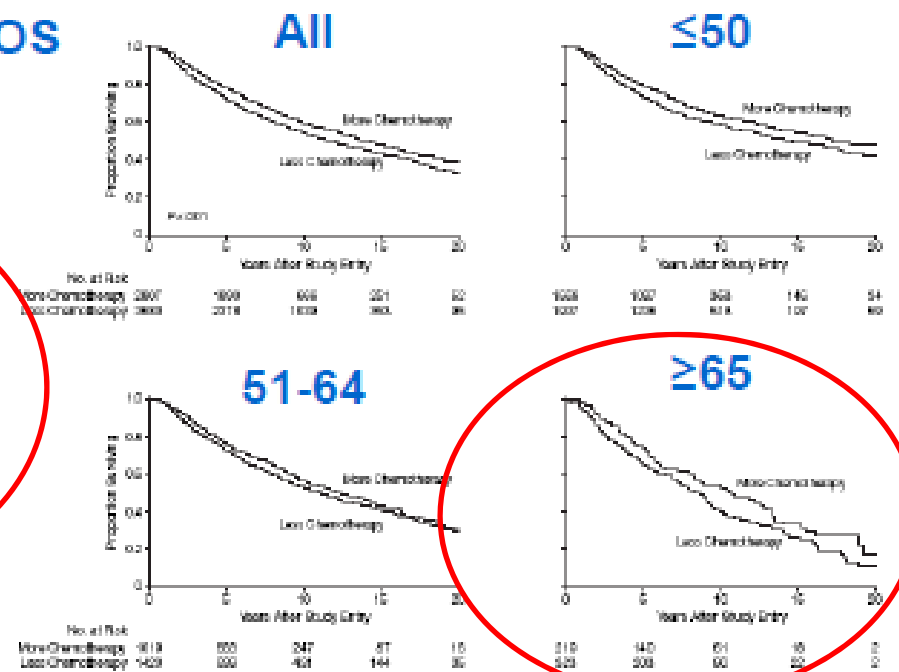
• **Results**

- Benefit **identical**
- Toxicity **careful!!**
- Toxic deaths 1.5%

- CALGB (1975-1999)
- 4 randomized trials
- 6487 pts

> 65 yo 542 (8%)  
> 70 yo 159 (2%)

OS



**Muss JAMA 2005**

# Adjuvant chemotherapy and mortality

		Giordano*	Elkin
	No. total No. w/CT	I-III, $\forall$ ER, 65+ 41,390 4,500	I-III, ER-, 66+ 5,081 1,711
pN	ER	HR (95% IC)	HR (95% IC)
pN0	$\forall$	1.05 (0.85-1.31)	NA
pN+	+	1.05 (0.85-1.31)	NA
both	-	NA	0.85 (0.77-0.95)
pN+	-	0.72 (0.54-0.96)	0.76 (0.65-0.88)
pN+ > 70 yo	-	0.74 (0.56-0.97)	

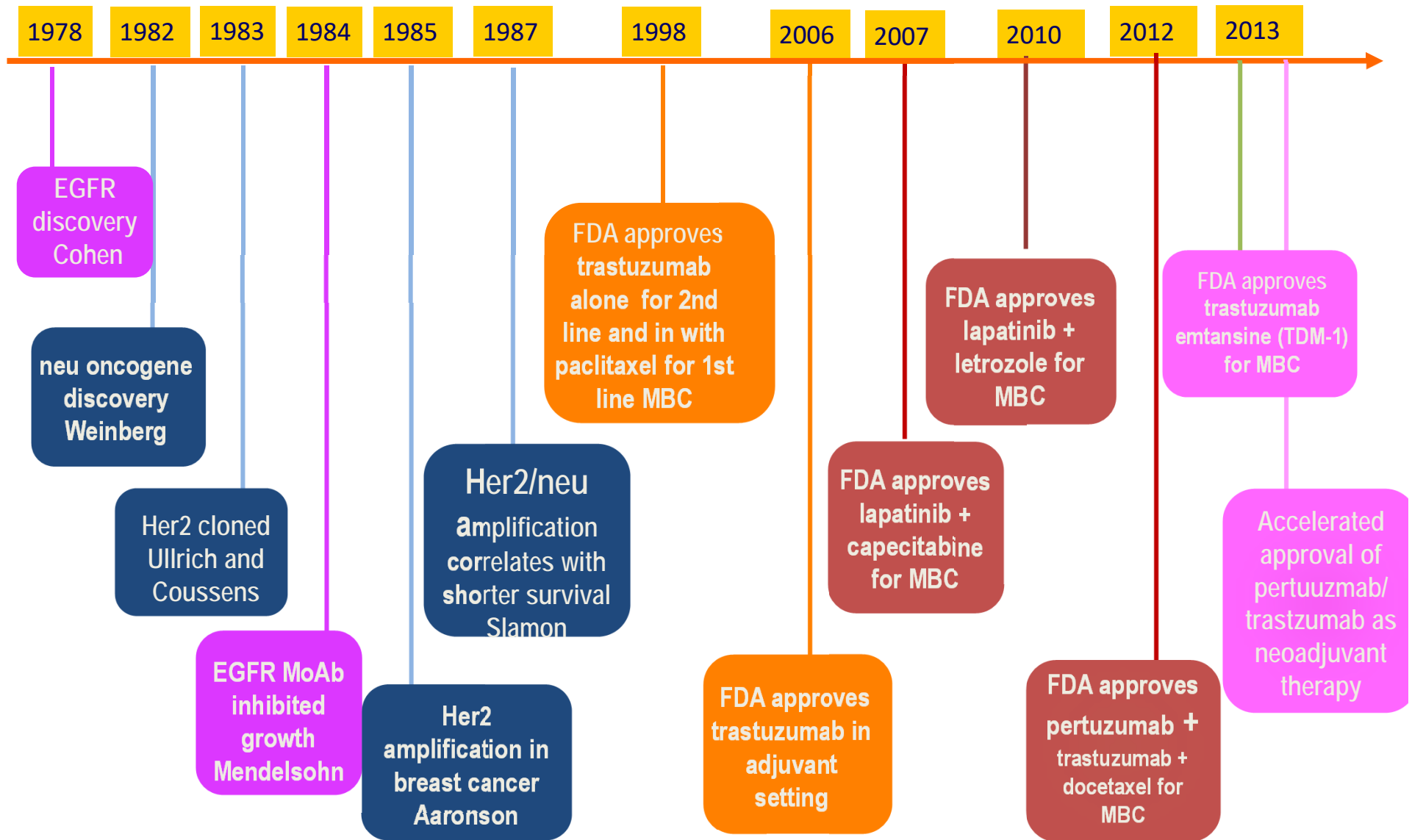
\*: BC specific mortality



**Adjuvant chemo is useful FIRST  
in ER-, pN0 or pN+, even > 70 yo**

Giordano & Elkin J Clin Oncol 2006

# Milestones of HER2/anti-HER2 therapies in BC



MBC : metastatic breast cancer; MoAb : monoclonal antibody

# DFS & OS w/ trastuzumab 1 year

Study	Follow-up (years)	N	DFS		OS	
			HR	p value	HR	p value
<b>HERA<sup>1-4</sup></b> CT+/-RT→H vs. CT+/-RT	1	3387	0.54	< 0.0001	0.76	0.26
	2	3401	0.64	< 0.0001	0.66	0.0115
	4	3401	0.76	< 0.0001	0.85	0.1087
	8	3401	0.76	< 0.0001	0.76	0.0005
<b>NCCTG N9831/ NSABP B-31<sup>5-7</sup></b> AC→TH→H vs. AC→T	2	3351	0.48	< 0.0001	-	-
	4	4045	0.52	< 0.001	0.61	< 0.001
	8.4	4046	0.60	< 0.0001	0.63	< 0.0001
<b>BCIRG 006<sup>8</sup></b> AC→TH→H vs. AC→T TCH vs. AC→T	5.5	3222	0.64	< 0.001	0.63	< 0.001
			0.75	0.04	0.77	0.04



**AMM FDA/EMA 2006**

CT, chemotherapy; DFS, disease-free survival; H, trastuzumab;  
HR, hazard ratio; OS, overall survival; RT, radiotherapy; T, taxane.

1. Piccart-Gebhart MJ, *et al*; *N Engl J Med* 2005; **353**:1659-1672;
2. Smith I, *et al*; *Lancet* 2007; **369**:29-36;
3. Gianni L, *et al*; *Lancet Oncol* 2011; **12**:236-244;
4. Goldhirsch A, *et al*; *Lancet* 2013 [Epub ahead of print];
5. Romond EH, *et al*; *N Engl J Med* 2005; **353**:1673-1684;
6. Perez EA, *et al*; *J Clin Oncol* 2011; **29**:3366-3373;
7. Romond EH, *et al*; SABCs 2012 (abstract S5-5; oral presentation);
8. Slamon D, *et al*; *N Engl J Med* 2011; **365**:1273-1283.



# Trastuzumab adjuvant & DFS

	HR all	(95%CI)	HR 60+	(95%CI)
HERA	0.64	0.54-0.76	0.91	0.59-1.41
NSABP-B31/N9831	0.48	0.39-0.59	0.41	0.24-0.68
BCIRG 006	0.61	0.37-0.65	NR	NR
FinHER	0.42	0.21-0.83	NR	NR
PACS-04	0.86	0.61-1.22	NR	NR

**> 60 yo ≤ 16% in HERA for ex!**

# *Elders with Breast Cancer Tend to Delay Seeking Medical Care and Present with a Later Cancer Stage*

Janice Tsang, Polly Cheung, Hang-mei Lee, Gary Tse,  
Sam Choy, Lorna Wong, Maria Shiu, Thomas Yau, Chun-Chung Yau

Steering Committee  
Hong Kong Breast Cancer Registry  
Hong Kong Breast Cancer Foundation

3<sup>rd</sup> March, 2017

Symposium on Elderly Primary Breast Cancer Women  
East Midlands Conference Centre, Nottingham

# Subjects and Methodology

- **13,265 female** patients with breast cancer, diagnosed between 2006 and 2015, from the Hong Kong Breast Cancer Registry were studied. Among them, **861 patients were aged 70 years or above.**
- **Chi square test was used to test for any significant differences between the elderly patients and patients of all ages in the following areas:**
  - How the breast cancer was first detected
  - Tumour characteristics
  - Types of treatment
- **Comorbidity of elderly patients were also assessed by using the **Charlson Comorbidity Index (CCI)****

# Elderly patients received more mastectomies & less chemotherapy and radiotherapy

	$\geq 70$ yo patients N (%)	Patients of all ages N (%)	P value
<b>Surgery</b>			
No	40 (5)	186 (2)	
Breast-conserving surgery (BCS)	91 (12)	3,785 (33)	<0.001*
Mastectomy (MTX)	608 (82)	6,562 (57)	
<b>Chemotherapy</b>			
I	8 (3)	1,544 (39)	<0.001*
II	30 (10)	4,035 (84)	<0.001*
III	32 (33)	1,574 (94)	<0.001*
IV	11 (42)	259 (87)	<0.001*
Targeted therapy	15 (14)	1,146 (55)	<0.001*
Endocrine therapy	576 (79)	8,605 (76)	0.098
<b>Radiotherapy</b>			
Among patients with BCS	80 (91)	3,591 (97)	0.010*
Among patients with MTX	198 (33)	3,654 (50)	<0.001*

\*p<0.05 indicates significant difference

# Elderly patients with higher CCI received more conservative treatment

	CCI =0 N (%)	CCI =1-2 N (%)	CCI ≥3 N (%)	P Value
<b>Surgery</b>				
No	48 (8)	8 (5)	11 (30)	<0.001*
<b>Chemotherapy</b>				
I	6 (4)	1 (2)	1 (25)	N/A
II	25 (10)	4 (7)	0 (0)	N/A
III	25 (34)	8 (38)	0 (0)	N/A
IV	--	--	10 (40)	N/A
Targeted therapy	12 (16)	3 (13)	0 (0)	N/A
Endocrine therapy	464 (75)	120 (77)	27 (77)	0.740
<b>Radiotherapy</b>				
Among patients with BCS	81 (88)	15 (88)	3 (60)	N/A
Among patients with MTX	146 (31)	42 (33)	9 (45)	N/A

\*p<0.05 indicates significant difference

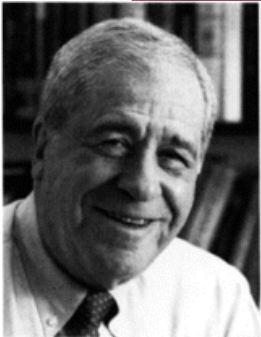
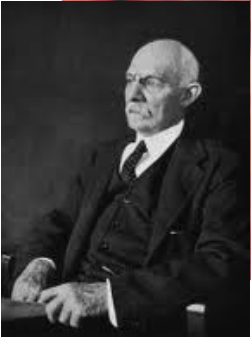


## Outline

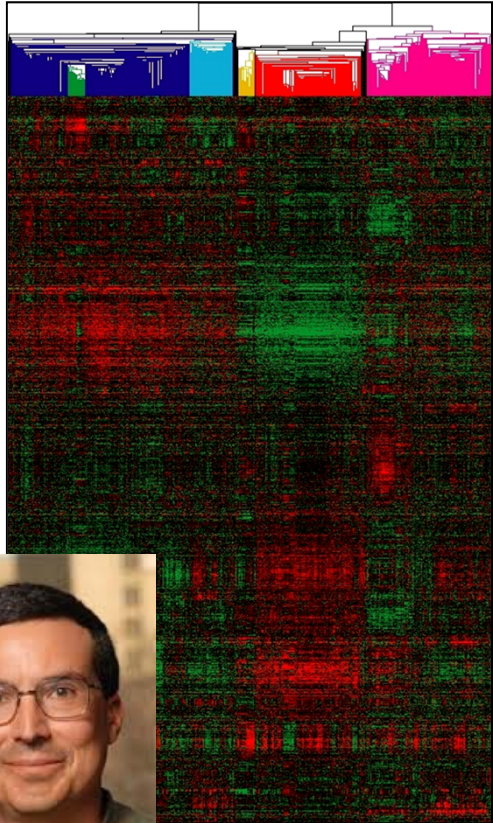
- The undeniable emerging needs
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- **Opportunities & Challenges**
- Conclusion



# Changing Portraits of Breast Cancer over the past decades...



claudin low  
Lum A Lum B Basal Her2



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## Found in translation: Prioritising research questions in breast cancer

22 Nov 2007

The key priorities that will impact on the future treatment of breast cancer have been identified by a group of experts on the disease. Research published in the online open access journal *Breast Cancer Research* may focus research resources onto the issues highlighted as top priorities.

A team led by Professor Mitch Dowsett, Head of Biochemistry at The Cancer Research UK Centre for Translational Research, based in London and Surrey, has led an international, web-based consultation on priorities for translational research. Translational research - the bridge between the lab to the clinic - holds huge promise for patients.

In this study, a database of over 40 clinicians, research scientists, academic and industry experts, at major breast cancer conferences, on-line and in print, were asked to register online and then log the most important research questions to tackle.

A steering committee reduced the 409 questions registered to 70 unique issues, from which participants were asked to vote for their 'top six'. In all, 420 participants from 48 countries voted; around half of voters classed themselves as clinicians.

The top research priority found was the identification of molecular signatures to select patients who could be spared chemotherapy. The second most pressing issue also involved chemotherapy, namely the identification of features to help clinicians choose the optimal chemotherapy regimen for individual patients.

While translational research in breast cancer has increased greatly over recent years, individual projects often reflect the immediate interests of the research group, rather than attempting to answer a specific question with potential to alter patient management. Identifying issues deemed important by the research community could help focus translational research resources, ensuring that opportunities for important clinical advances aren't missed.

"This appears to be a novel way to identify the most important challenges for improving breast cancer treatment and prevention" explains Professor Dowsett. "The work will allow investigators globally to select the most relevant clinical research questions in their efforts to translate the major advances in basic science to improvements in the clinical management of this common malignancy. I am grateful to the participants from 48 countries who made this possible."

Article:

[A web-based consultation on priorities for translational research in breast cancer](#)  
Mitch Dowsett, Aron Goldhirsch, Alan Coates, Josep Serra, William Wood and Giuseppe Viale  
*Breast Cancer Research* 2007, 9:R81 (22 November 2007)

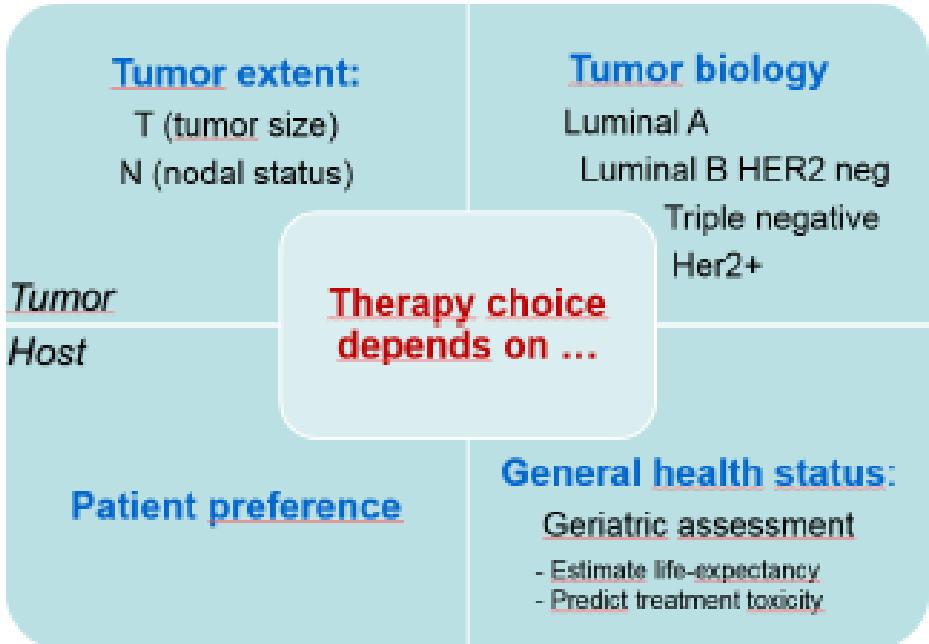
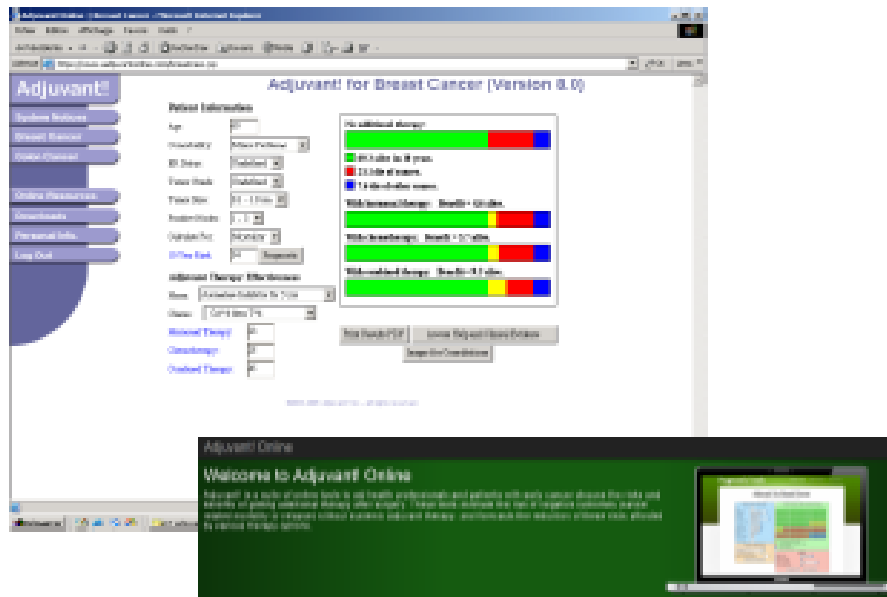
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The top research priority found was “the identification of molecular signatures to select patients who could be spared from chemotherapy. The second most pressing issue also involved chemotherapy, namely the identification of features to help clinicians choose the most optimal chemotherapy regimen for individual patients”

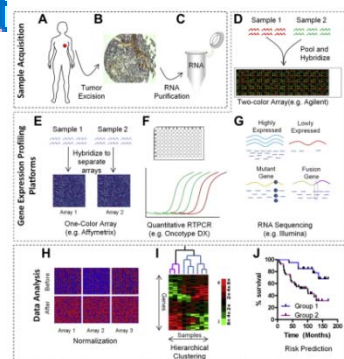


# Standard decision tools



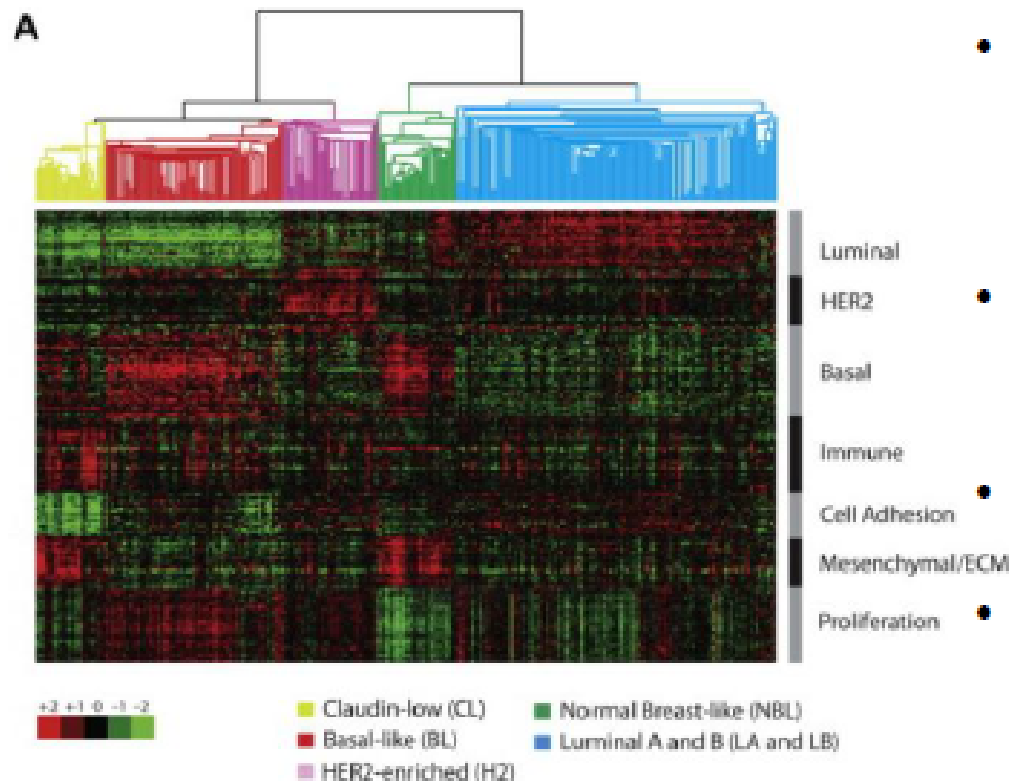
- **Adjuvant! Online**
- **Predict**

not accurate in older patients  
quite accurate for OS prediction



De Glas Lancet Oncol 2014 & Br J Cancer 2016

# Early 2000s: 1<sup>st</sup> GEP (intrinsic classification)



- Quantification of mRNA or cDNA of genes involved in tumour proliferation
- To identify patients requiring chemo despite good standard prognostic factors
- To avoid chemo in others
- Better individual risk stratification

Prat Mol Oncol 2011

# Various options of molecular genomic profiling tools...

	MammaPrint	Oncotype DX	Breast Cancer Index	Maquant DX	PAM 50 ROR	EndoPredict
<b>Provider</b>	Agendia	Genomic Health	Biotheranostics	Ipsogen	NanoString	Sividon
<b>Type of Assay</b>	70-gene assay	21-gene recurrence score	2-gene ratio (M/I) and molecular grade index	Genomic grade	50-gene assay	12-gene assay
<b>Type of Sample</b>	Fresh or frozen or FFPE	FFPE	FFPE	Fresh or frozen or FFPE	FFPE	FFPE
<b>Technique</b>	DNA microarray or qRT-PCR	qRT-PCR	qRT-PCR	DNA microarray or qRT-PCR	qRT-PCR	qRT-PCR
<b>Clinical Application</b>	Prognosis of NO, < 5 cm, stage I/II, age < 61	Prediction of recurrence risk in ER+ and NO treated with TAM	Prognostic in ER+, prediction of response to TAM	Molecular grading for ER+, histologic grade II disease	Originally for intrinsic subtyping, recurrence prediction	Recurrence prediction for ER+ HER2-
<b>Results Presentation</b>	Dichotomous, good or poor prognosis	Continuous variable	Continuous variable	Dichotomous, GGI I or GGI III	Continuous variable	Dichotomous, low or high risk
<b>Level of Evidence</b>	I	I	III	III	I	I
<b>FDA Approval</b>	YES	NO	NO	NO	YES	NO

Abbreviations: ER+, estrogen receptor-positive; FDA, U.S. Food and Drug Administration; FFPE, formalin-fixed, paraffin-embedded; GGI, Genomic Grade Index; qRT-PCR, quantitative reverse transcription polymerase chain reaction; TAM, tamoxifen.

## MammaPrint

Summary of Results

PATIENT NAME: Last Name, First Name **doe, Jane Doe** REPORTED DATE: **21-Feb-2014**

ADJUVANT RESPONSE TO THERAPY

Summary of Results: **High Risk Luminal-type (B)**

Risk of Recurrence: **High Risk**

Molecular subtype: **Luminal-type**

MammaPrint® PPMF, 70-Gene Breast Cancer Recurrence Assay

**Risk of distant recurrence risk @ 5 years w/ no treatment**

PATIENT NAME: **Jane Doe-Jane Doe-Jane** REPORTED DATE: **21-Feb-2014**

ADJUVANT RESPONSE TO THERAPY

Distant Metastasis Free Survival @ 5 yrs for MammaPrint High Risk?

Endocrine: 76%  
Endocrine + Chemotherapy: 98%

General results

**General results**

## OncotypeDX

Genomic Health | oncotypedx

Breast Cancer Report - Node Negative

PROGNOSTIC SCORE: **6**

10-Year Risk of Distant Recurrence: **5%**

10-Year Risk of Distant Recurrence after 5 Years of Tam, Based on the Recurrence Score Result (From NSABP B-14)

**Risk of distant recurrence @ 10 years w/ TAM 5 years**

Genomic Health | oncotypedx

Breast Cancer Report - Node Positive

PROGNOSTIC SCORE: **6**

10-Year Risk of Distant Recurrence: **11%**

10-Year Risk of Distant Recurrence after 5 Years of Tam, Based on the Recurrence Score Result

**Risk of distant recurrence @ 5 years w/ TAM 5 years**

Personal results



**MINDACT**  
 Microarray In Node-negative  
 and 1 to 3 positive lymph node  
 Disease may Avoid ChemoTherapy  
 EORTC 10041 / BIG 3-04



The NEW ENGLAND  
 JOURNAL of MEDICINE

ESTABLISHED IN 1912

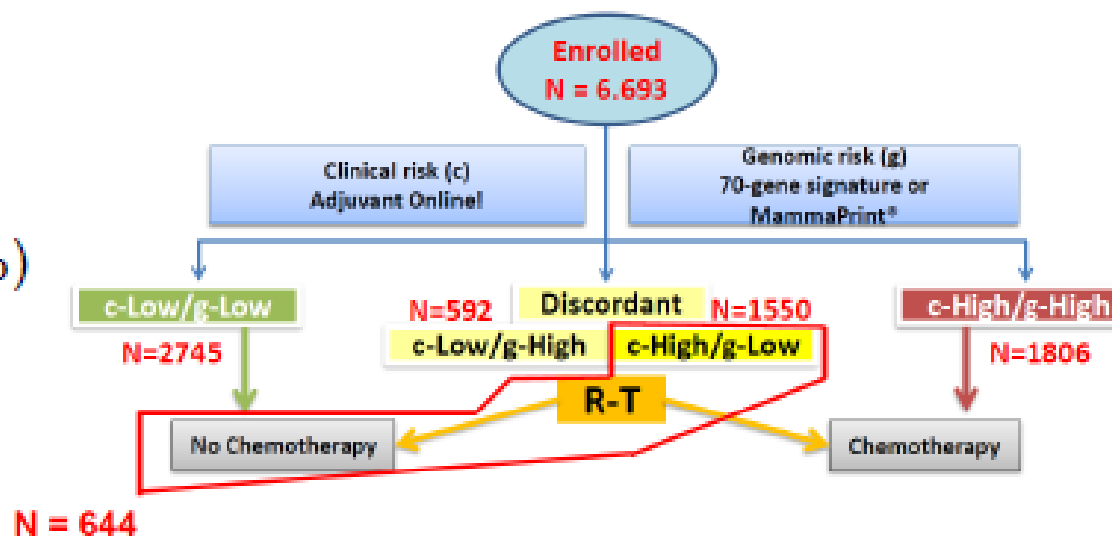
AUGUST 23, 2010

VOL. 362 NO. 8

70-Gene Signature as an Aid to Treatment Decisions  
 in Early-Stage Breast Cancer

F. Cardoso, L.J. van't Veer, J. Bogaerts, L. Slaets, G. Viale, S. Delalogo, J.-Y. Pignatelli, E. Braña, S. Casaretto, M. DeLorenzi, A.M. Glas, V. Gollanopoulos, T. Goulioti, S. Konec, E. Matos, B. Meulemans, P.A. Neljehuis, U. Nitz, R. Passalacqua, P. Ravdin, I.T. Rubio, M. Saghatelyan, T.J. Smilg, C. Sotiriou, L. Stork, C. Striebel, G. Thomas, A.M. Thompson, J.M. van der Hoven, P. Vajsbek, R. Bernardi, K. Tsyferkalis, E. Rutgers, and M. Piccart, for the MINDACT Investigators\*

- 6,600 pts < 70
  - FEB 2007-AUG 2011
  - 11,291 registered pts
  - 6,673 enrolled (59.1%)



# People in their eighties...



## OUTCOME DISPARITIES BY AGE AND 21-GENE RECURRENCE SCORE RESULT IN HORMONE RECEPTOR-POSITIVE (HR+) BREAST CANCER

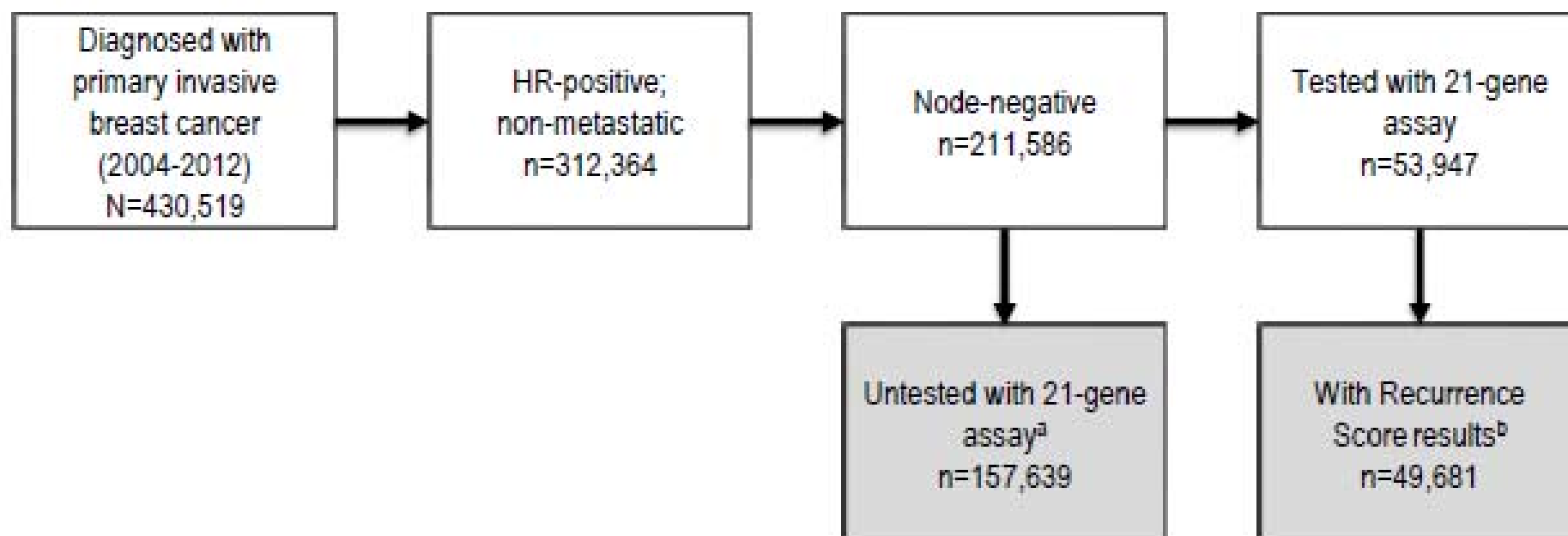
Shak S,<sup>1</sup> Miller DP,<sup>1</sup> Howlader N,<sup>2</sup> Gliner N,<sup>1</sup> Howe W,<sup>3</sup> Schussler N,<sup>3</sup> Cronin K,<sup>2</sup> Baehner FL,<sup>1,4</sup> Penberthy L,<sup>2</sup> Petkov VI<sup>2</sup>

1. Genomic Health, Inc., Redwood City, CA, USA
2. National Cancer Institute, Rockville, MD, USA
3. IMS, Inc., Calverton, MD, USA
4. University of California, San Francisco, San Francisco, CA, USA

### Methods

- SEER demographics, tumor characteristics, reported CT use, and BCSM available through 2013
- Genomic Health provided RS electronically to SEER, per registry operations
- Analysis population: N0, HR+ (by SEER and RT-PCR), HER2-negative (by RT-PCR), diagnosed between January 2004 and December 2012
  - Excluded: N+, prior invasive tumors, or concurrent multiple tumors
- RS groups standard cutpoints (18, 31)
- Actuarial estimates of survival (cause-specific and overall) and BCSM computed through 5 years with 95% CI
- The log-rank test was used to compare the three RS groups

# SEER Population - STROBE Diagram



a. Untested cohort without RS results includes patients with HER2+ breast cancer because HER2 status was not reported to SEER before 2010.

b. Tested cohort with RS results excludes patients with HER2+ breast cancer, based on 21-gene assay quantitative single-gene HER2 result. Median follow-up for younger (<70 years) and older (≥70 years) patients were 45 and 40 months, respectively.

# Patient Testing and Demographics

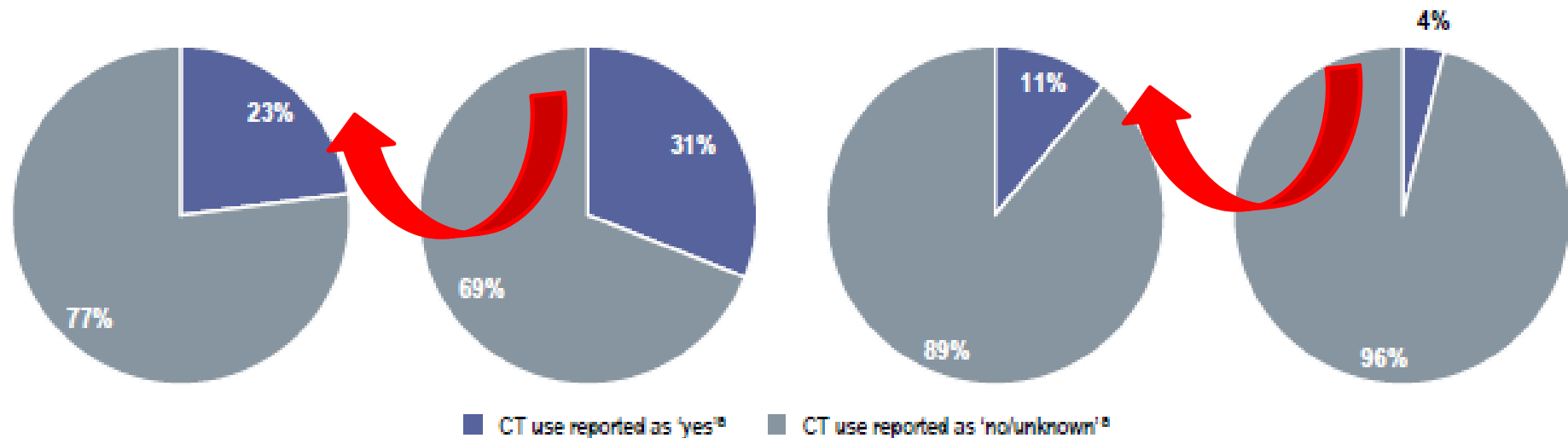
		Age <70 years		Age ≥70 years	
		Tested (N=43,693)	Not Tested (N=100,519)	Tested (N=5,988)	Not Tested (N=57,120)
		%	%	%	%
Sex <sup>a</sup>	Female	99	99	99	99
Race	White	84	81	87	87
	Black	8	9	7	7
	Asian or Pacific Islander	8	10	5	6
	Am. Indian/Alaska Native	<1	<1	<1	<1
Socioeconomic status, quintile	Lowest SES	11	13	15	14
	Second lowest SES	15	17	17	18
	Middle SES	19	20	21	21
	Second highest SES	23	23	23	22
	Highest SES	32	28	25	25

- Almost 6,000 patients ≥70 years with RS results
- Testing occurred 3.2 times less frequently in patients ≥70 years compared to <70 years
- Testing rates were similar by race and socioeconomic status



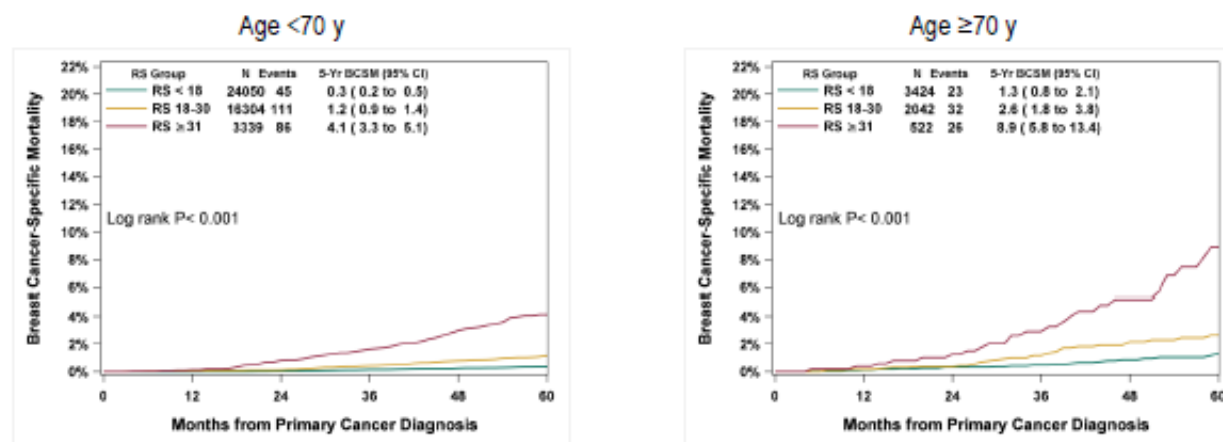
# Reported Chemotherapy (CT) Use

Age <70 years		Age ≥70 years	
Tested (N=43,693)	Not Tested (N=100,519)	Tested (N=5,988)	Not Tested (N=57,120)



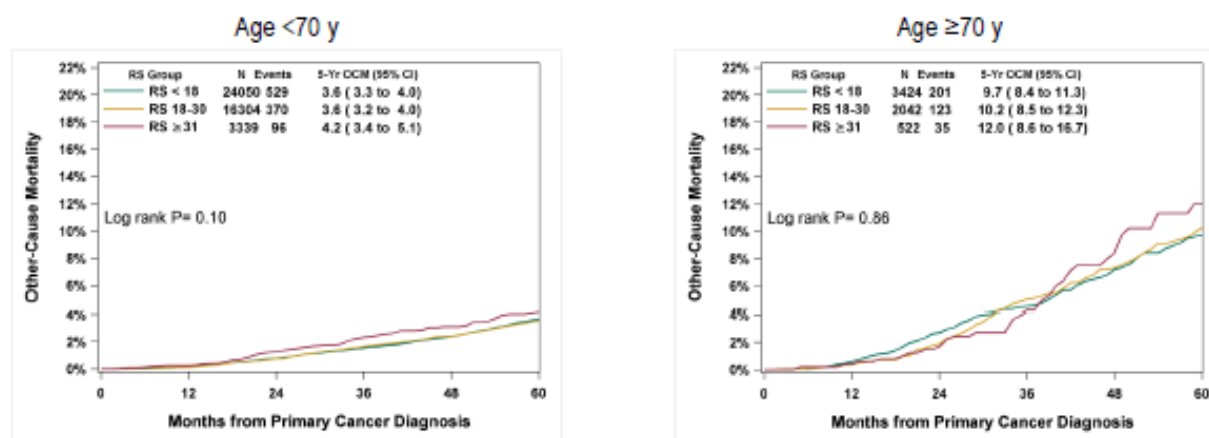
- CT use was lower in patients ≥70 years, in both RS-tested and untested cohorts

## 5-year BCSM by Age and RS Group



- RS predicts BCSM in both age groups ( $p < 0.001$ )
- Low 5-y BCSM was observed with RS <18 in both age groups
- Higher 5-y BCSM was observed with RS 18-30 and RS  $\geq 31$  in older patients

## 5-year Other-Cause Mortality by Age and RS Group



- As expected, RS group does not predict other-cause mortality ( $p = \text{NS}$ )
- As expected, higher other-cause mortality was observed in older patients

## 5-year BCSM (95% CI) by Age in Tested and Untested Patients

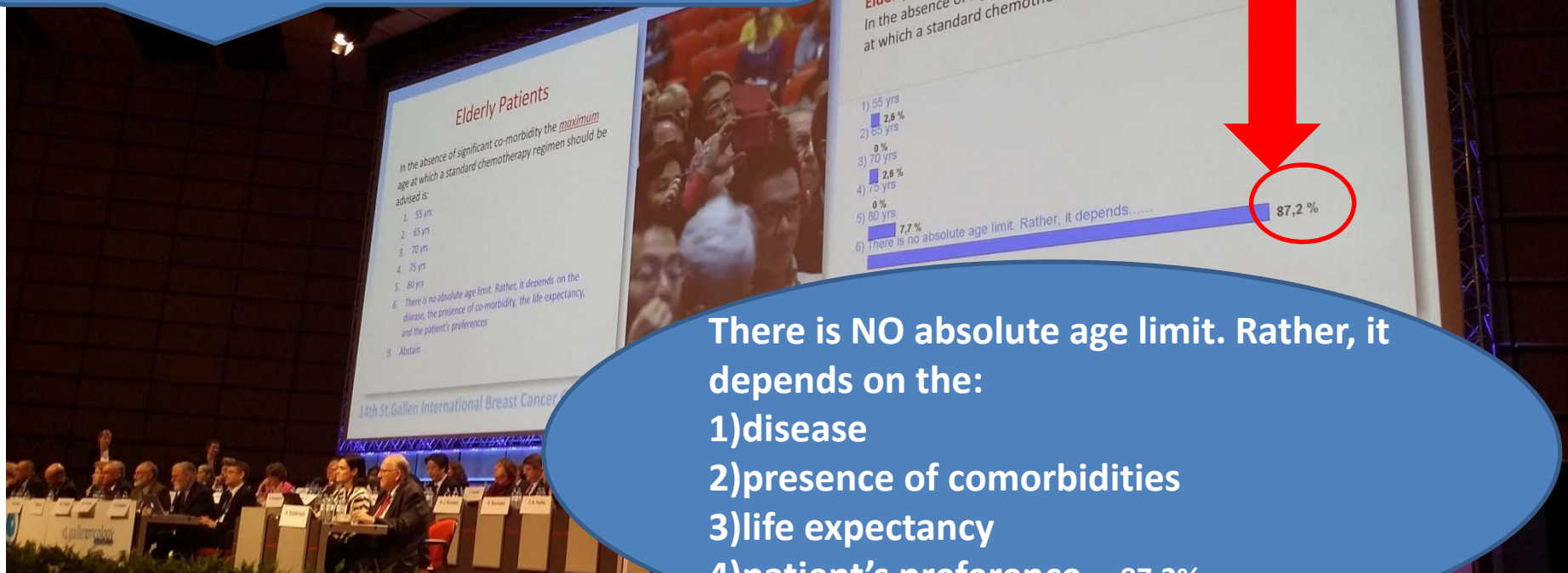
	RS < 18			RS 18-30			RS ≥31			Untested		
	N	CT Use <sup>a</sup> (% of N)	5-y BCSM (95% CI)	N	CT Use <sup>a</sup> (% of N)	5-y BCSM (95% CI)	N	CT Use <sup>a</sup> (% of N)	5-y BCSM (95% CI)	N	CT Use <sup>a</sup> (% of N)	5-y BCSM (95% CI)
<70 y	24050	7%	0.3% (0.2%, 0.5%)	16304	37%	1.2% (0.9%, 1.4%)	3339	73%	4.1% (3.3%, 5.1%)	100519	31%	2.3% (2.2%, 2.4%)
≥70 y	3424	2%	1.3% (0.8%, 2.1%)	2042	14%	2.6% (1.8%, 3.8%)	522	52%	8.9% (5.8%, 13.4%)	57120	4%	5.5% (5.2%, 5.7%)
70-74 y	2116	2%	1.1% (0.6%, 2.0%)	1245	17%	2.4% (1.4%, 3.9%)	320	61%	6.2% (3.2%, 11.9%)	17647	8%	2.8% (2.6%, 3.2%)
75-79 y	968	2%	1.9% (0.9%, 4.0%)	590	11%	2.4% (1.1%, 5.2%)	142	43%	11.6% (5.5%, 23.8%)	16445	4%	4.3% (4.0%, 4.7%)
≥80 y	340	1%	1.0% (0.2%, 4.5%)	207	6%	4.8% (2.3%, 9.9%)	60	32%	20.5% (9.6%, 40.6%)	23028	2%	8.6% (8.2%, 9.1%)

- Notably, 5-y BCSM is relatively high in untested patients at all ages; this deserves further study



# St. Gallen Breast Cancer Conference 21<sup>st</sup> March, 2015, Vienna.

In the absence of significant co-morbidity the  
MAXIMUM age at which a standard  
chemotherapy regimen should be advised is...



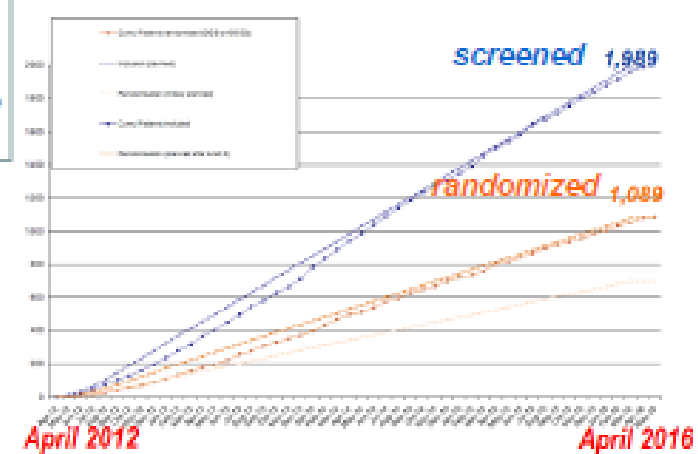
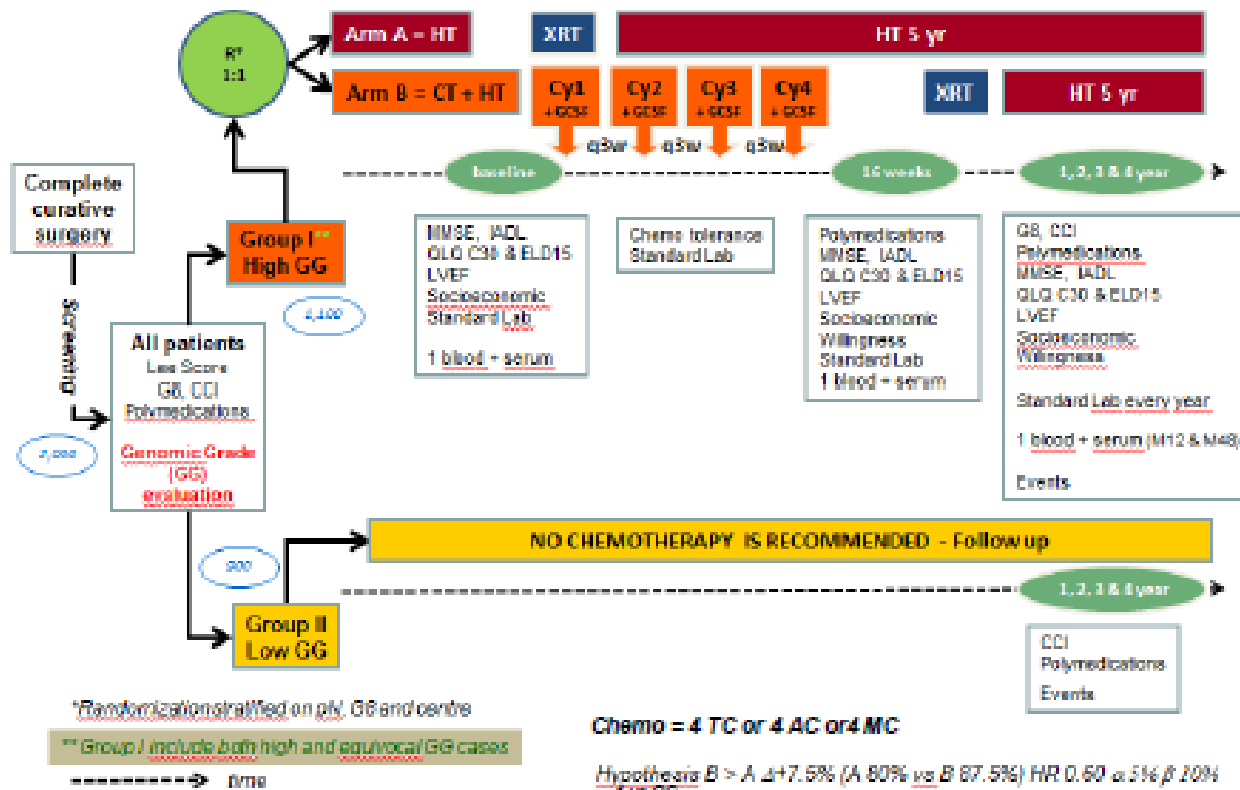
There is NO absolute age limit. Rather, it depends on the:

- 1) disease
- 2) presence of comorbidities
- 3) life expectancy
- 4) patient's preference ... 87.2%

# Adjuvant chemotherapy

ER+ (ongoing study)

## ASTER 70s (EUDRACT N° 2011-004744-22, PHRC national 2011, NCT01564056)



ADVERTISEMENT



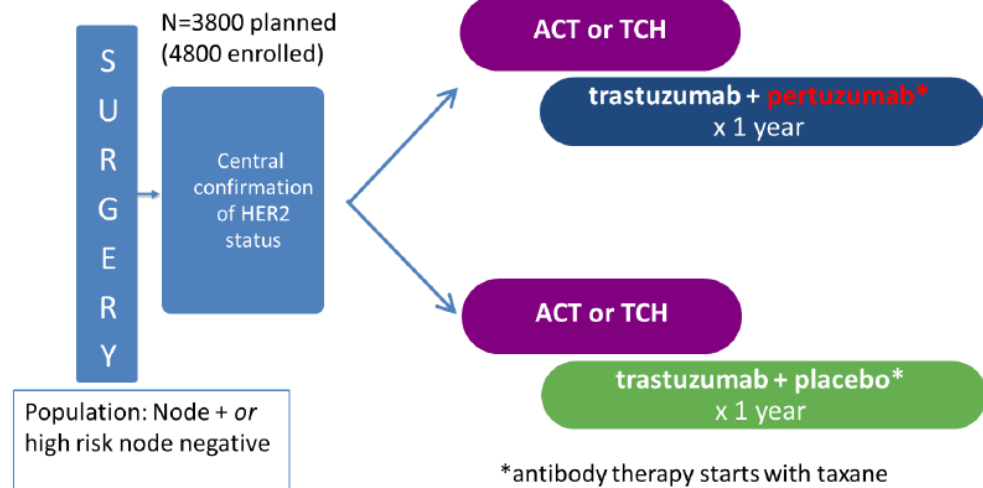
Providing Oncology Professionals the Resources They Trust

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# Phase III APHINITY Study: Adjuvant Pertuzumab/Trastuzumab/Chemotherapy Increased Invasive Disease-Free Survival in HER2-Positive Breast Cancer

By The ASCO Post  
 Posted: 3/2/2017 10:55:14 AM  
 Last Updated: 3/2/2017 10:55:14 AM

## APHINITY



A=doxorubicin, E=epirubicin, C=cyclophosphamide, T=taxane (paclitaxel or docetaxel), F=5-fluorouracil, H=trastuzumab, P=pertuzumab



## Outline

- The undeniable emerging needs
- Current Dilemmas for Elderly BC patients
- Opportunities & Challenges
- **Conclusion**



# Our Future Directions...

## Young Patients

“Quantity of life” – to strive to live longer

Family and social obligations

## Oncologist’s perspective

Investigations and Treatment

Response and Toxicities

- RECIST

- NCI CTC V 4.0

- Survival (DFS, PFS, OS)

- “Fast-Moving” world

“Molecular Portrait”

## GEP

- Identifying individual patient who can be spared or benefitted from chemo (systemic therapy)

## Elderly Patients

Quality of life +++

Independent, and staying at home

## Geriatrician’s perspective

Symptoms and Diagnosis

Quality of Survival

- Amount of life with good QoL

- cognition

- functional status

- QoL

- nutrition

“Global Portrait” – aging population

## CGA

- Identifying individual elderly patient who will benefit from systemic therapy



# Our Future Directions...

## Young Patients

“Quantity of life” – to strive to live longer

Family and social obligations

## Oncologist’s perspective

Investigations and Treatment

Response and Toxicities

- RECIST

- NCI CTC V 4.0

- Survival (DFS, PFS, OS)

- “Fast-Moving” world

“Molecular Portrait”

## GEP

- Identifying individual patient who is not harmed or spared or benefitted from chemotherapy (systemic therapy)

## Elderly Patients

Quality of life +++

Preference for staying at home

## Geriatric perspective

Accurate Diagnosis

Prognosis

Survival with good QoL

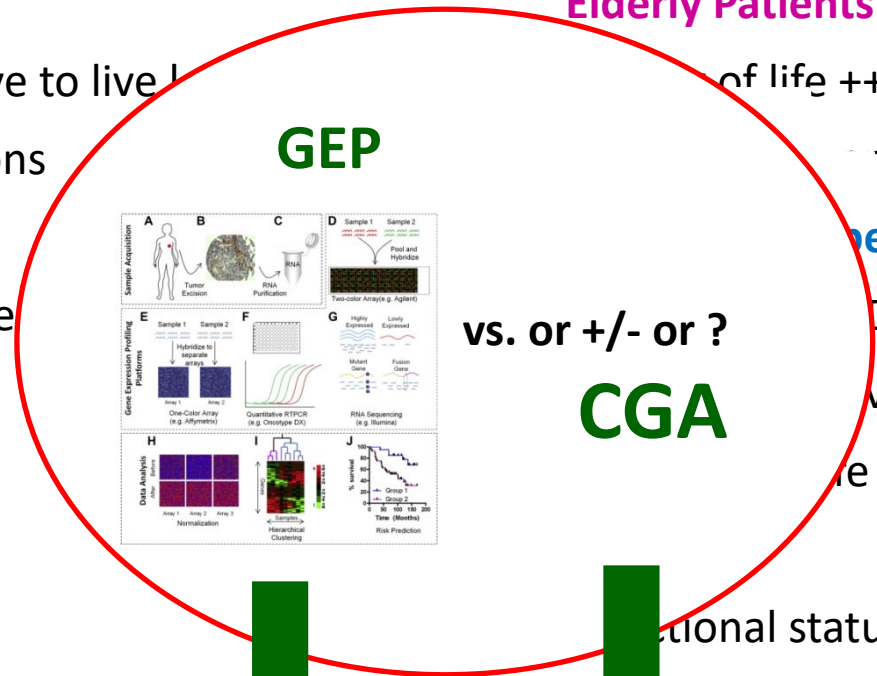
Functional status

- Cognitive function

- Medication

“Clinical Portrait” – aging population

- Identifying individual elderly patient who will not benefit from systemic therapy



**Genomic Defect Targeted Therapy**

**CGA Defect Targeted Geriatric Intervention**

FEC, AACR, FAC, ASCO, anti-PDL1, anti-PD1, CMF, SABCS, PD-1, PDL1, DXR, PK/PD, CEX, 5FU CDDP, Calvert AUC, ESMO, Chatelut AUC, CTC, TILs, population PK, EORTC, FOLFIRI, ctDNA, FOLFOX 7, CPA, DFS, CALGB, DDFS, OS, TTP, NCI, CYP P450, JCO, JNCI, HER2, PI3K, mTOR, Phase 0, ECCO, ib and ab, Unicancer, EORTC, SWOG, CALGB, etc.

Charlson, CIRSG, CGA, AD, MCI, MNA, GDS, MMS, ADL, IADL, GFI, CMR2, JAGS, EUGMS, G8, CARG, Oncodage, VES-13, TRFs, JGO, NIA, SoFOG, Walter's score, Lee's score, CRASH, etc.



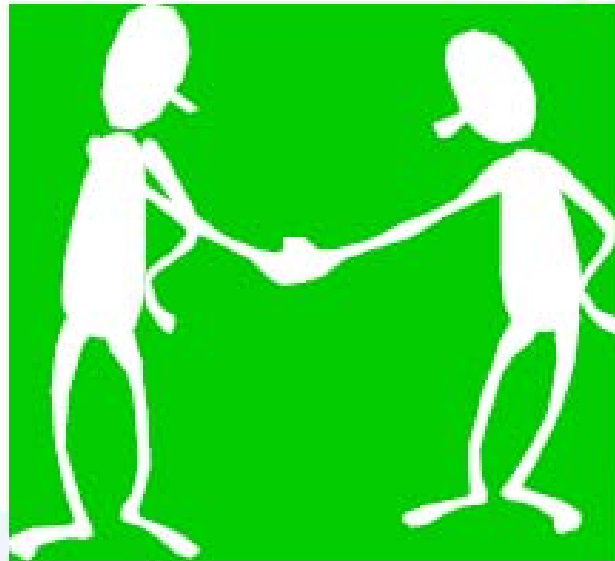
**Oncologist**



**Geriatrician**



**SIOG**  
INTERNATIONAL SOCIETY  
OF GERIATRIC ONCOLOGY



FEC, FAC, SoFOD, ADL, IADL, CME, SABCS, DXR, PK/PD, CEX, G8, EORTC, 5FU CDDP, MCI, Calvert and Chatelut AUC, CARG, GDS, population PK, AD, FOLFIRI, MMS, FOLFOX, CPA, CRASH, SWOG, DFS, OS, TTP, NCI, GERICO, TILs, CARG, anti-PDL1, anti-PD1, EORTC TFE, JCO, JNCI, Charlson, JGO, CIRSG, PD-1, PDL-1, ctDNA, EGS, EGA, MNA, GFI, Unicancer, Lee's score, JAGS, etc.

**To be practice changing,  
let us be practice sharing!**

**Geriatric Oncology Collaboration**

**From Professor Etienne Brain, Immediate Past President of the SIOG**

# Acknowledgements

The GBCC 2017 OC members  
 The University of Hong Kong  
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 The Hong Kong Breast Cancer Foundation  
 The International Society of Geriatric Oncology (SIOG)  
 All the breast and non-breast elderly patients and collaborators



香港乳癌基金會策動  
 A Hong Kong Breast Cancer Foundation Initiative

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9-11 NOV.

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**SAVE THE DATE - November 9-11, 2017**  
**[www.siog.org](http://www.siog.org)**



To improve the care of older patients with cancer around the world



# Elderly Breast Cancer – Systemic Therapy

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**Founding Convenor**

**Hong Kong Breast Oncology Group**

**Scientific Committee Meeting**

**International Society of Geriatric Oncology (SIOG 2017)**



**20<sup>th</sup> April, 2017**

