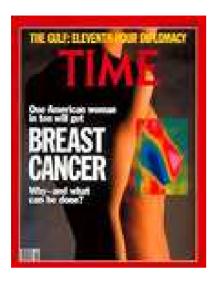
Breast Cancer Disparity in Asian Women

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Department of Health, Behavior, & Society



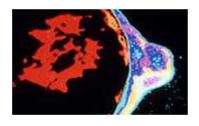
2009 Global Breast Cancer Conference Seoul, Korea October 8-10, 2009























Topics to be covered

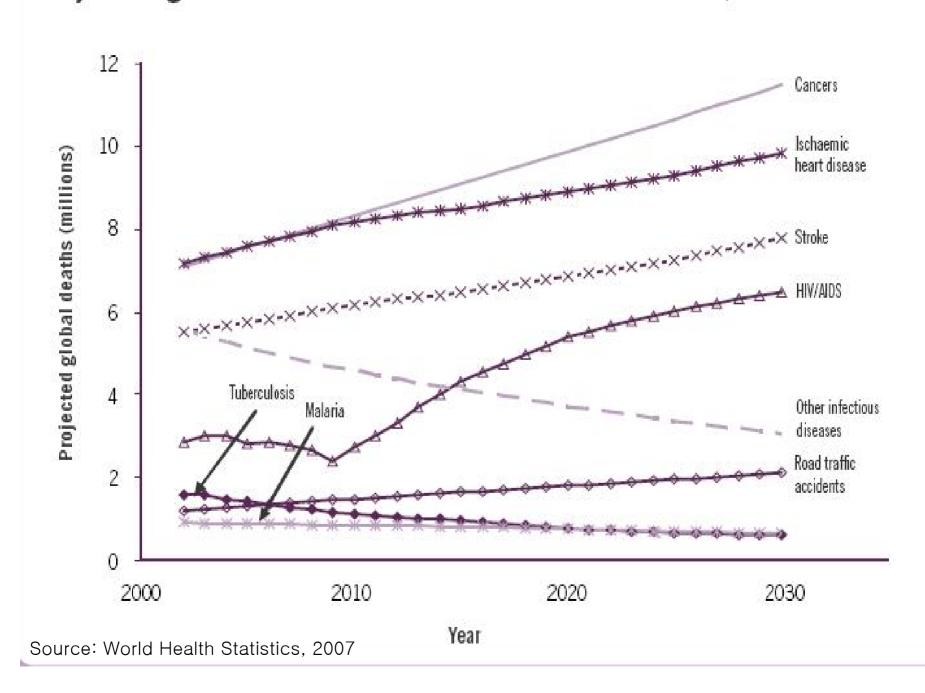
Breast cancer burdens in Asia

Hereditary Breast Cancer in Asia

Breast Cancer Control in Asia

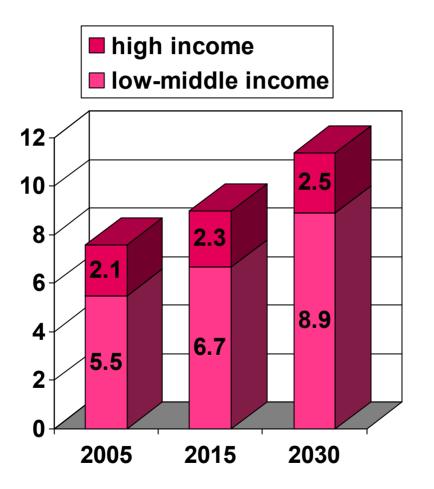
Future directions

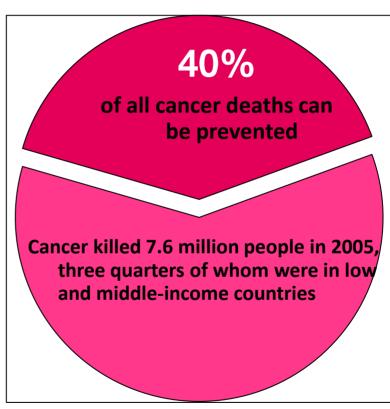
Projected global deaths for selected causes of death, 2002-2030¹⁵



Projections of Cancer Mortality

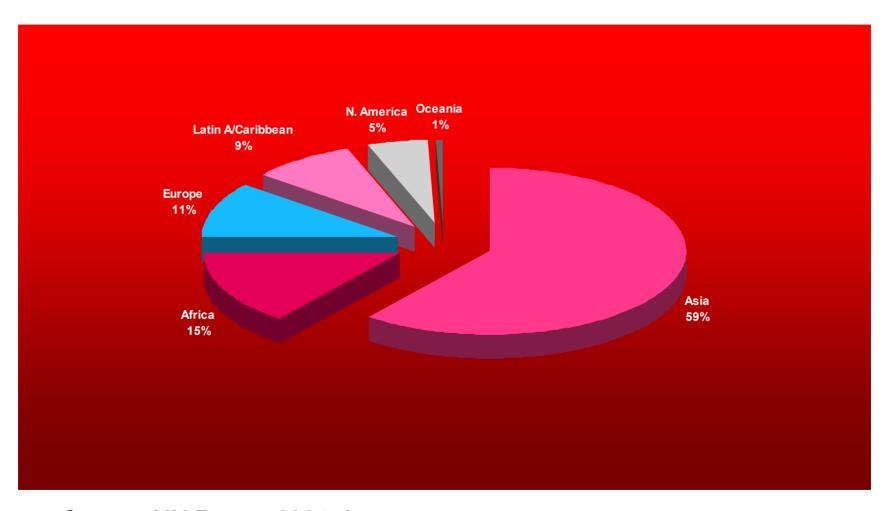
WHO projects worldwide





World Population in 2008

(total: 6,707 millions)



Source: UN Report 2004 data

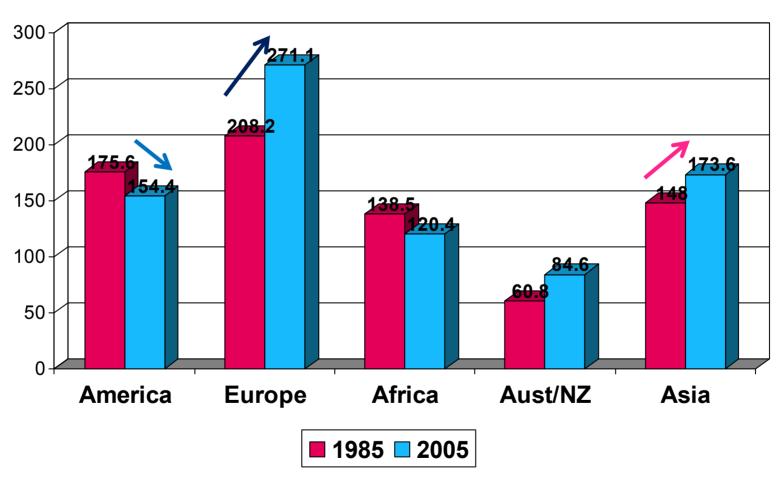
Breast Cancer Statistics

Worldwide

- Breast cancer is the most common diagnosed cancer among women.
- More than 1.1 million women are newly diagnosed with breast cancer annually.
 - This represents about 10% of all new cancer cases and 23% of all females cancers
- With more than 410,000 deaths each year, breast cancer accounts for about 14% of all female cancer deaths.
- An estimated 4.4 million breast cancer patients diagnosed within the last five years.

UICC: Breast Health Global Initiative

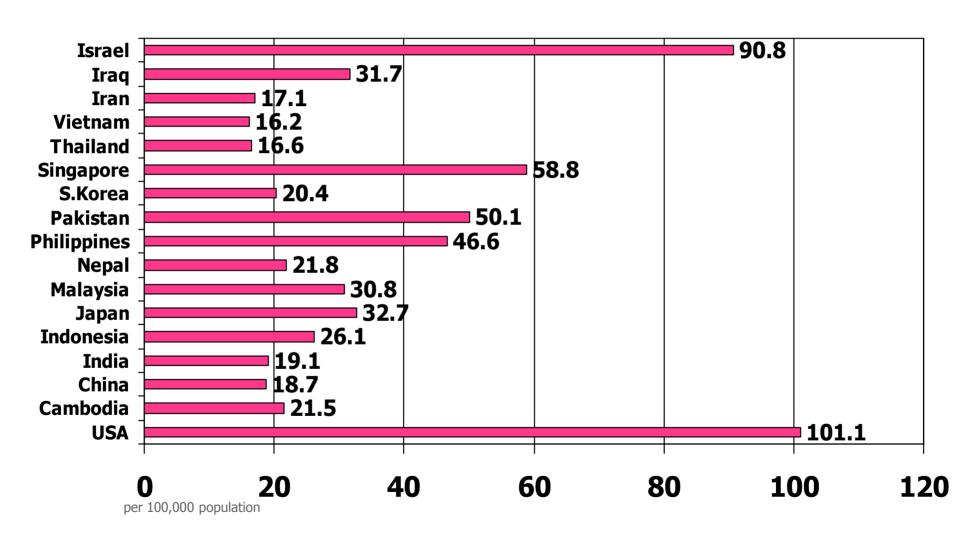
Comparison of age-standardized breast cancer incidence, 1985 to 2005



Parkin et al. (1993). Estimates of worldwide incidence of eighteen major cancers in 1985. Int J Cancer, 54: 594-606 Parkin et al. (2005). Global Cancer Statistics, 2002. CA Cancer J Clin, 55,74-108

Age-standardized breast cancer incidence, 2002

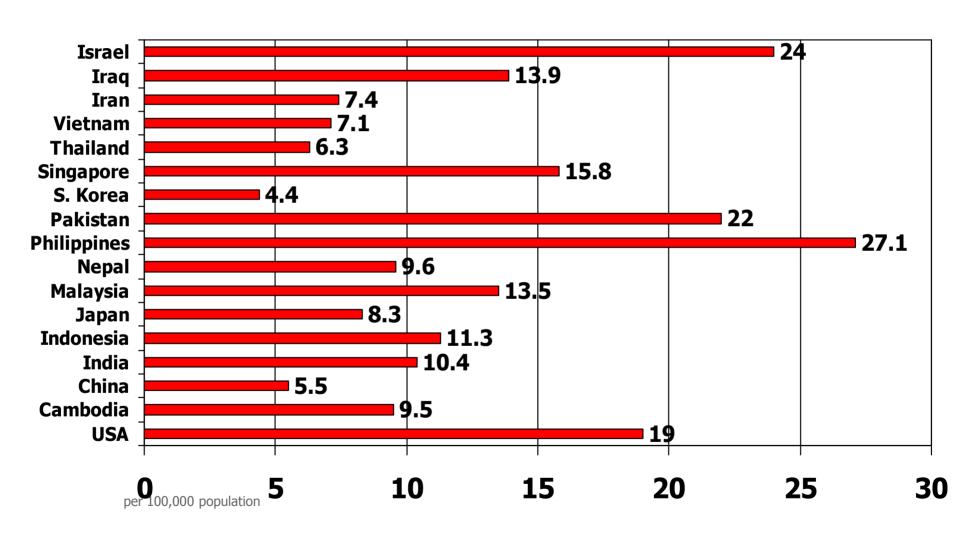
Selected countries of Asia



Source: International Agency for Research on Cancer, 2007

Age-standardized breast cancer mortality, 2002

Selected countries of Asia



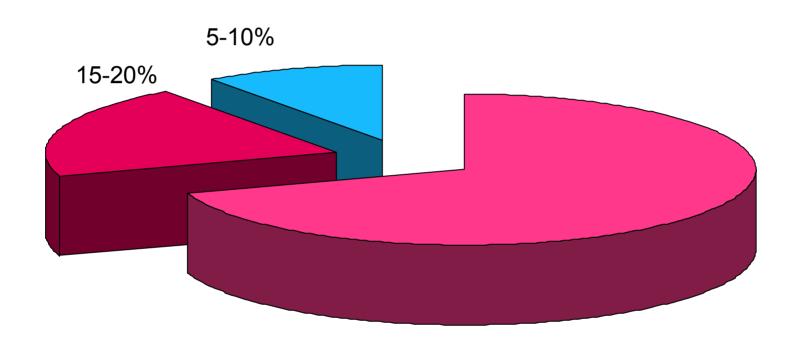
Source: WHO Statistical Information System (WHOSIS), 2007

Breast Cancer Disparity in Asia

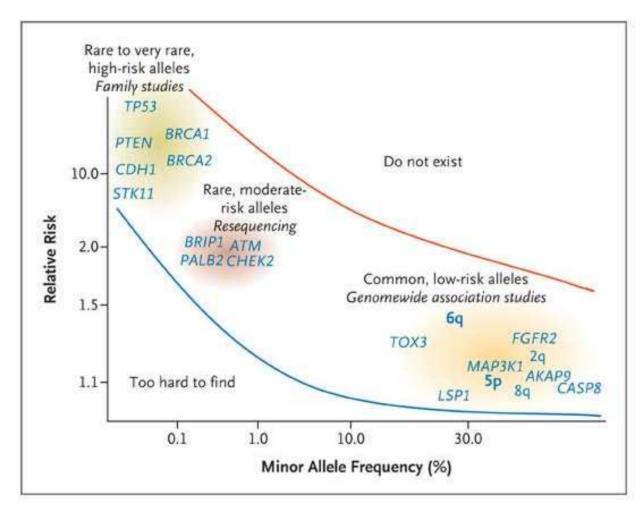
- By 2020, 70% of all breast cancer cases worldwide will be in developing countries (IARC 2002)
- Low breast cancer incidence but high breast cancer mortality
- Early age-of-onset
 - Increased incidence in young age group in Asia
 - Mean age is around 50 years, and prevalent age group is 40-49 years old

BREAST CANCER GENETICS

How much breast cancer is hereditary?



■ Sporadic ■ Family clusters ■ Hereditary



Foulkes, WN. (2008). Inherited susceptibility to common cancers. The New England Journal of Medicine, 359, 20, 2143-2153



Causes of hereditary susceptibility to breast cancer

Gene

Contribution to hereditary Breast Caner

| BRCA1 | ~30% | ~50% |
|--------------------|------|------|
| BRCA2 | ~20% | |
| TP53 | <1% | |
| PTEN | <1% | |
| Undiscovered genes | ~50% | |

BRCA1 & BRCA2

BRCA1

- Tumor suppressor gene
 on chromosome 17
- Autosomal dominant transmission
- Protein has role in genomic stability
- ~500 different mutation reported
- 50%-85% lifetime risk of breast cancer (often early age onset)

BRCA2

- Tumor suppressor gene
 on chromosome 13
- Autosomal dominant transmission
- Protein has role in genomic stability
- ~300 different mutation reported
- 50%-85% lifetime risk of breast cancer

Prevalence of BRCA mutations

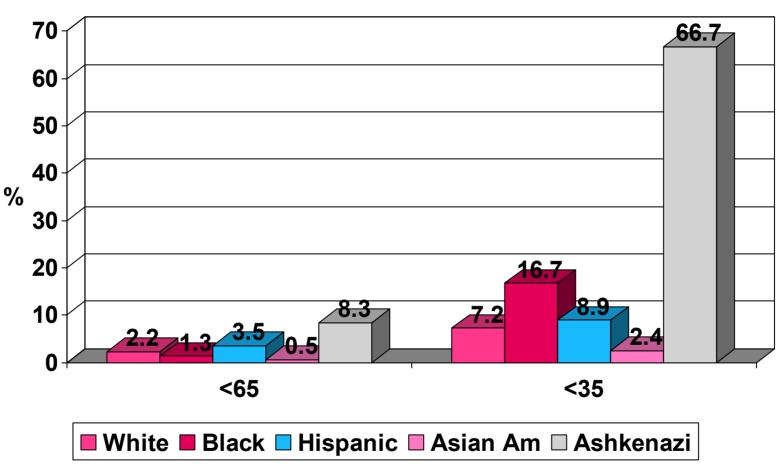
- 2-4% Breast Cancer due to BRCA1/BRCA2 in N.
 America, Europe, Israel, Australia (all ages)
 - Malone et al., 2006; Newman et al., 1998; John et al., 2007
- Higher prevalence of BRCA mutations for younger age (<35 years old): 3.5% to 7.2%
- Data on BRCA mutations in Asian patients with breast cancer are sparse.
 - Prevalence of BRCA1 among Chinese early onset breast cancer: 7 to 8 % (Sng et al., 2000; Ho et al., 2000; Suter et al., 2004)

Prevalence of BRCA1 mutation by race

- Northern California Breast Cancer Family Registry, 1996-2005
- Population-based female breast cancer patients younger than 65 years old at diagnosis
- Estimate BRCA1 carrier prevalence by 5 racial/ethnic group in the US

Prevalence of BRCA1 mutation by ethnicity

(n=3,181)



Source: John et al. Prevalence of pathogenic BRCA1 mutations carriers in 5 US racial/ethnic groups. JAMA. 2007; 298(24): 2869-2876

Risk of breast cancer mutations underestimated for Asian women

- To test BRCA mutation prediction models in Asian Americans: BRCAPRO & Myriad II
- 200 Asian women and 200 matched whites
- Under-predicted the true number of clinically important mutations among Asians

Comparison

| | White | Asian American |
|----------|-------|---------------------|
| BRCAPRO | 24/25 | 25/49 |
| MyriadII | 25/25 | 26/49 |
| P-value | >.69 | <3x10 ₋₇ |

Source: Kurian et al. (2008). Performance of BRCA1/2 mutation prediction models. Journal of Clinical Oncology. 26(29), 4752-4758.

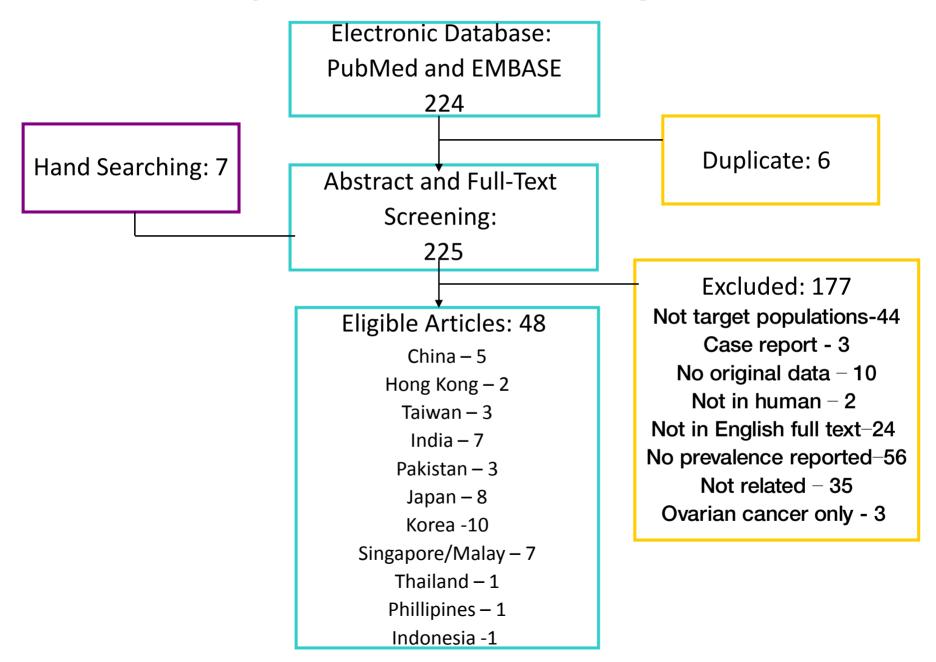
Prevalence of BRCA1 and BRCA2 Germline Mutations for Women with Breast Cancer in Asia

- We assessed the prevalence of BRCA1 and BRCA2 germline mutations in women with breast cancer in Asian countries.
- Systematic literature review of BRCA1 and BRCA2 in Asia from 8 studies (Liede & Narod, 2002)
- We extended this systematic review
- We conducted meta-analysis.

Literature Search

- Searched PubMed and EMBASE
 - English publications after 1990
 - Keyword search
 - BRCA1 or BRCA2
 - Asia
 - breast cancer
 - Focused on East, South, and Southeast Asian countries

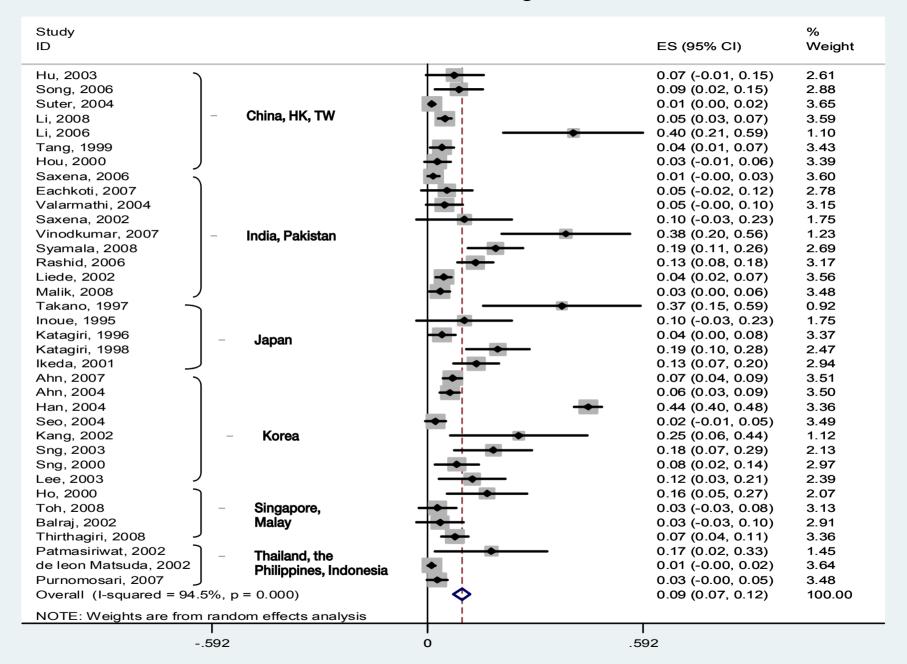
Summary of Search and Screening Process



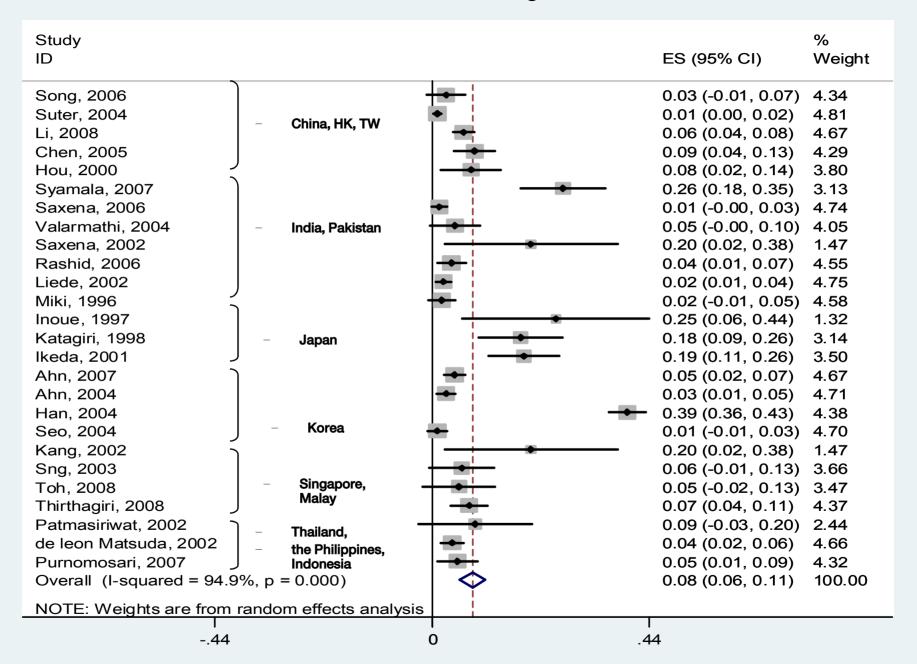
Meta-Analysis

- Extracted the prevalence data
 - Overall analysis by gene types (e.g., BRCA1, BRCA2)
 - Subgroup analysis
 - By age of early onset
 - By family history
- Random effect meta-analysis:
 - To get pooled prevalence of germline mutations for BRCA1, BRCA2 and BRCA1 and BRCA2
 - Used DerSimonian and Laird method
 - Used STATA10 (metan command)

Prevalence of BRCA1 Germline Mutations among Breast Cancer Women in Asia



Prevalence of BRCA2 Germline Mutations among Breast Cancer Women in Asia

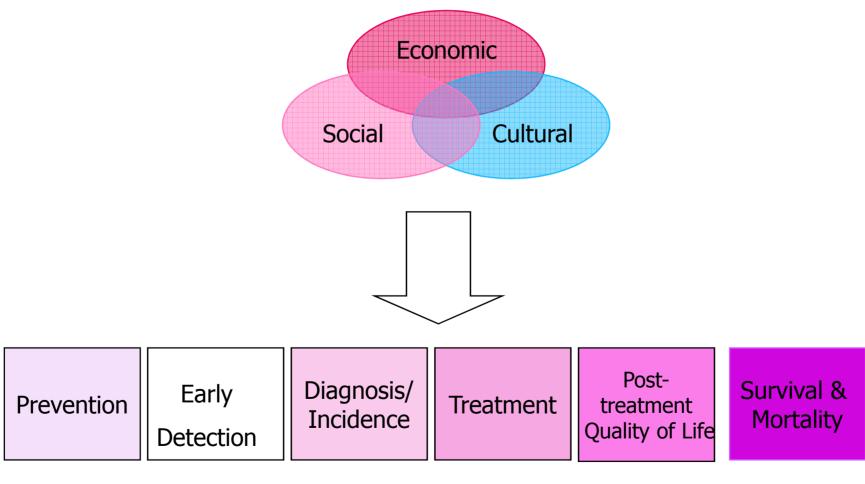


Summary

- Pooled prevalence of Asian population (8%-10%) is higher than U.S. population (2%-4%)
 - May be due to higher proportion of high risk population included in this review
 - More population-based studies are needed
- This BRCA mutation estimation is important for counselors and clinicians to provide most evidence-based and ethically sound approach to the women with high risk of breast cancer.

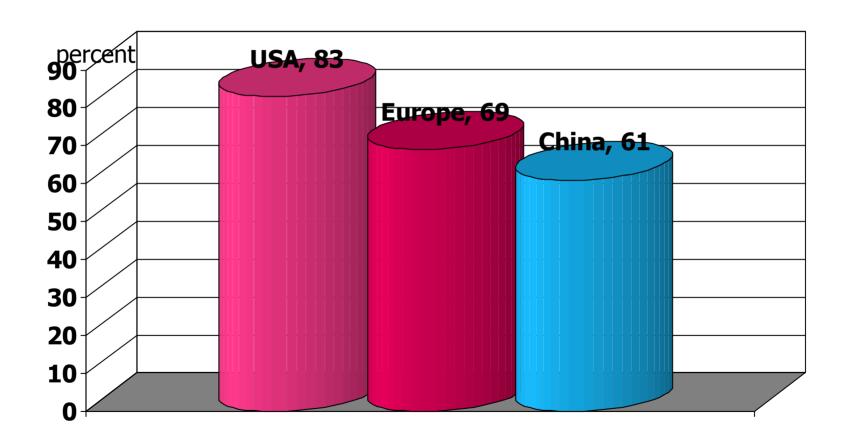
BREAST CANCER CONTROL

Comprehensive Breast Cancer Care



Source: Adapted Institute of Medicine

5-year breast cancer survival rates



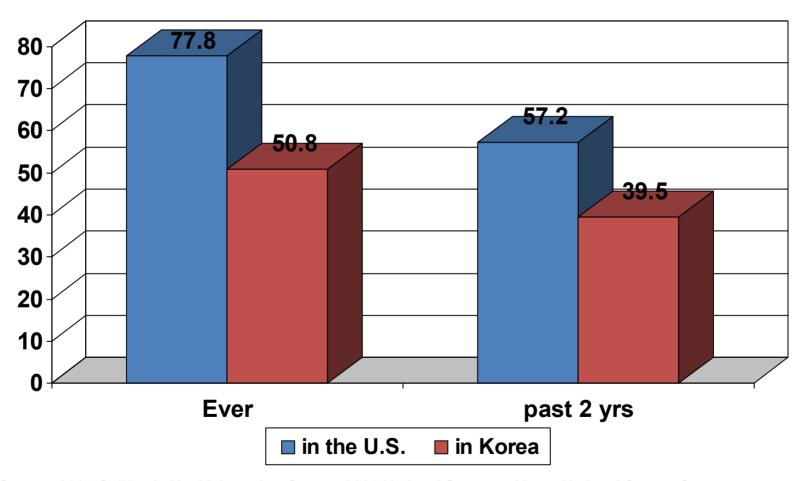
Source: International Agency for Research on Cancer, 1998

Stage of Breast Cancer Diagnosis in Asia

- In Asia (India, Malaysia, China, Iran), about 50 to 60% women diagnosed at stage 3 and 4 (late stage)
- In Japan, 90% diagnosed at stages 0 to 2 (early stage)
- In South Korea, 45.2% diagnosed at stages 0 to 1 (early stage).

Comparison of having a mammogram.

Korean American women and Korean women in Korea (40 years+)



Source: 2005 California Health Interview Survey; 2005 National Survey at Korea National Cancer Center Choi et al. (in press). Comparison of breast cancer screening rates between Korean American women and Korean women in Korea: what makes these differ? Journal of Women's Health

Challenges of Breast Cancer Control in Asia

- Low public awareness of breast cancer and BC prevention
- Social norm and taboo about cancer or breast
- Limited resources on breast health
- Low screening rates
- Late stage at diagnosis
- Lack of surveillance program for reliable data

Future Direction for Global Breast Cancer Control Strategy

- Systematic & comprehensive cancer control
 - Every country should have cancer control plan
- Surveillance program through cancer registry
 - To generate reliable data on the magnitude and patterns of breast cancer
- Prevention will play a pivotal role in the countries with limited resources
 - Best practices with limited resources



Thank you