



## SCIENCE AND EVIDENCE: EXPOSING THE MISINFORMATION USED TO CONFUSE THE IMPORTANCE OF BREAST CANCER SCREENING

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## Nothing to disclose



#### SHOULD WE AGREE TO DISAGREE?

Can't experts look at the same information and come to different conclusions?



## Experts Disagree ??

## The Earth certainly looks flat!!





## THERE ARE FACTS



## ALTERNATIVE FACTS

There have been more than 50 years of misinformation in the effort to reduce access to mammography screening



JUST IN CASE – A FEW FACTS

- 1. There is little if any "overdiagnosis" of invasive breast cancer.
- No one has ever seen an invasive breast cancer, detected by mammography, "disappear" on its own
- 3. Screening reduces the rate of advanced cancers



JUST IN CASE – A FEW FACTS

- 4. "False positives" is a misnomer. These are simply recalls from screening for a few extra pictures or an ultrasound.
- 5. The only "harm" that is affected by delaying screening is the recall rate.
- 6. Screening is the main reason that deaths have declined. Therapy saves lives when breast cancers are treated earlier.



Mammography screening is one of the major medical advances in the last 50 years. It has undergone greater scrutiny and more challenges than virtually any other medical intervention.

Opposition has persisted for over 50 years despite continually mounting evidence of benefit.



#### BREAST CANCER SCREENING ALTERNATIVE FACTS

Kopans DB. The Breast Cancer Screening "Arcade" and the "Whack-A-Mole" Efforts to Reduce Access to Screening. Semin Ultrasound CT MR. 2018 Feb;39(1):2-5.10.1053/j.sult.2017.06.002. Epub 2017 Jun 28.



#### BREAST CANCER SCREENING ALTERNATIVE FACTS

- There is no benefit from screening (1960-2009)
- 2. We can't possibly screen all women -(1970)
- 3. The radiation from the mammogram will cause more cancers than will be cured (1976)
- 4. There is no benefit from screening women ages 40-49 (1993)
- 5. The parameters of screening change abruptly at the age of 50 (1994-1997)



### **ALTERNATIVE FACTS**

- 6. Screening women in their forties should be based on their risk of developing breast cancer (2008).
  7. "Overdiagnosis" - Invasive cancers detected by mammography would "melt away" if not detected by screening (2009).
- 8. Improved therapy is the reason lives are being saved
- 9. Screening doesn't work because it did not reduce "all cause mortality" in the randomized controlled trials."



## BREAST CANCER SCREENING ALTERNATIVE FACTS

10. Screening doesn't work because it doesn't reduce the rate of advanced cancers.

11. Screening everyone is "old fashioned". We need "value based" screening.

(Kopans DB. The Breast Cancer Screening "Arcade" and the "Whack-A-Mole" Efforts to Reduce Access to
Screening. Semin Ultrasound CT MR. 2018 Feb;39(1):2-5.10.1053/j.sult.2017.06.002. Epub 2017 Jun 28.)



#### THE DECADES LONG EFFORT TO REDUCE ACCESS TO SCREENING

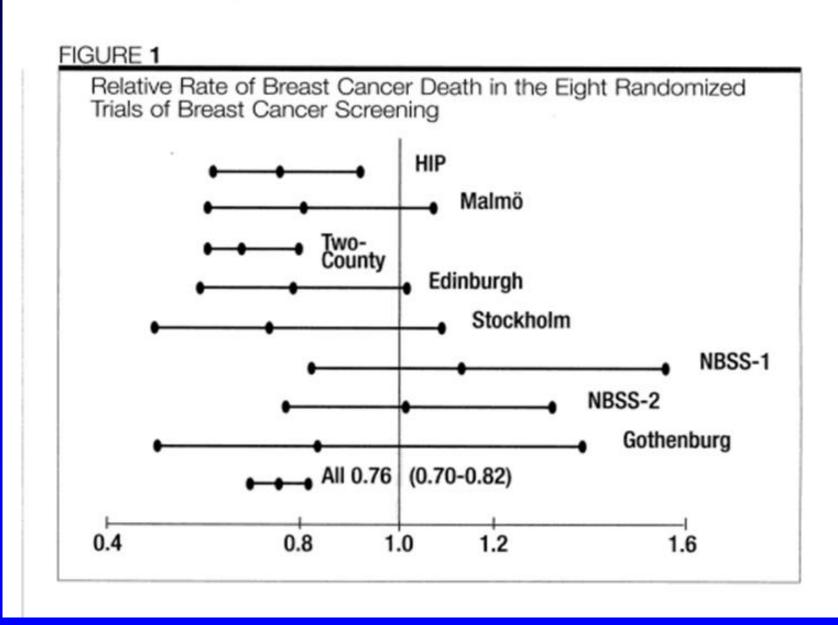
Science has shown, repeatedly, that screening reduces deaths from breast cancer, while specious arguments have been created, and then refuted in an effort to reduce access.



Randomized, controlled trials have, unequivocally, shown a statistically significant mortality reduction for screening for women ages 40-74

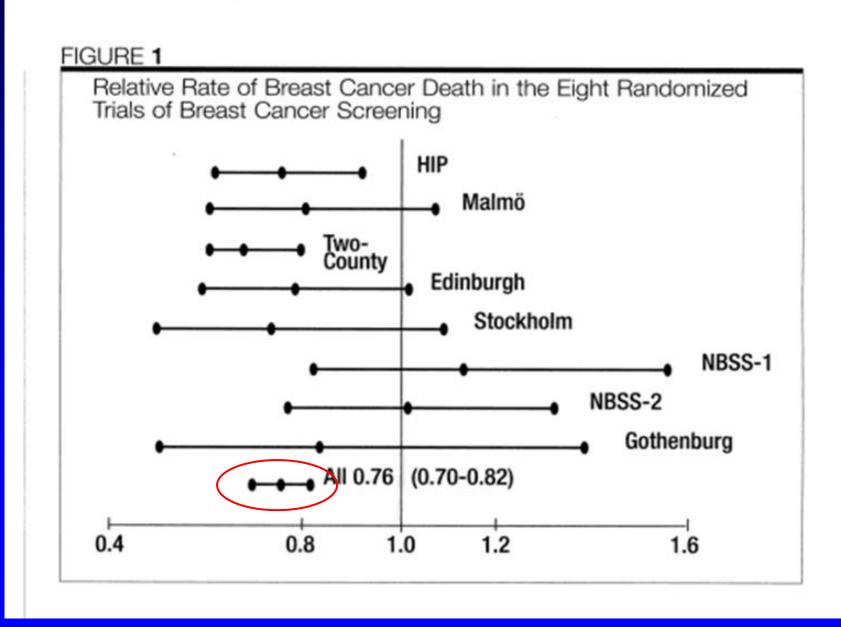


CA Cancer J Clin 2002;52:68-71



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CA Cancer J Clin 2002;52:68-71



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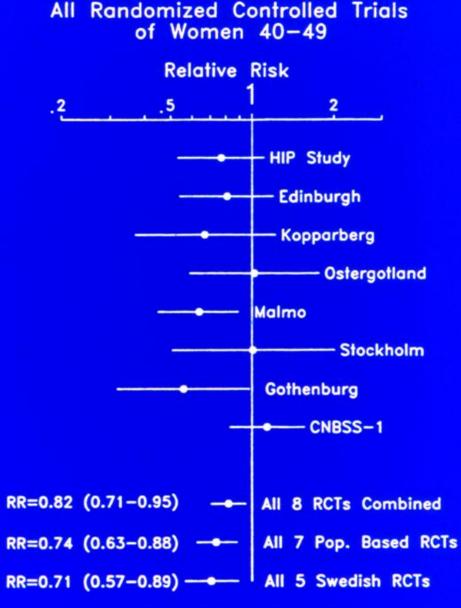
2000

Despite early and, scientifically, inappropriate analyses claiming no benefit, a statistically significant mortality reduction is clear for women ages 40-49.



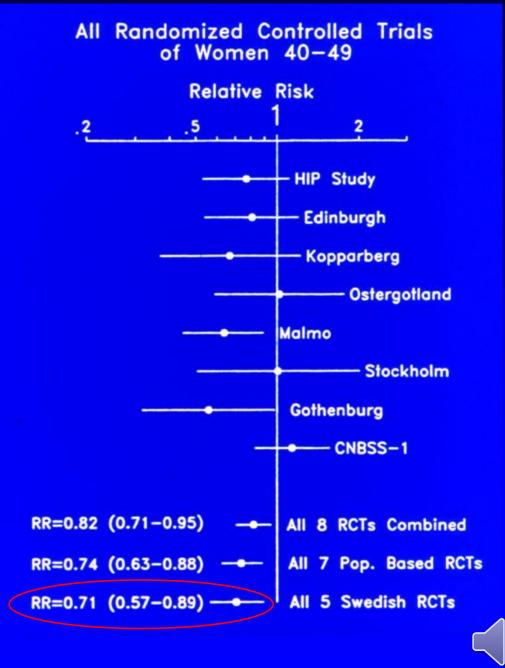
#### **SCREENING FOR WOMEN AGES 40-49**

Although the RCT were never intended to be analyzed by age groups, the data show a benefit from screening women ages 40-49. This was provided to, and ignored by the Panel at the 1997 Consensus Development Conference



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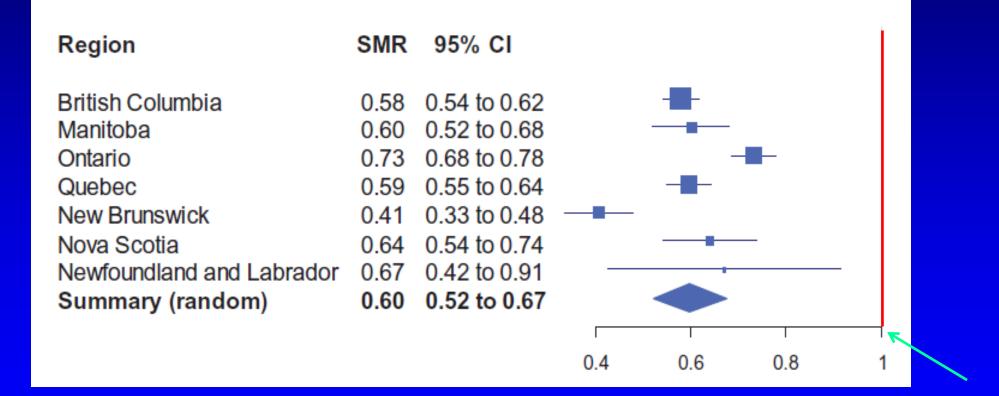


RCT'S actually underestimate benefit due to "non-compliance" and "contamination".

When screening is introduced into general populations the death rate from breast cancer declines to an even greater extent

#### SCREENING IN CANADA IS SAVING LIVES

Comparing women who participate in screening and those who do not, the death rate for the screened women is 40% (range 27%-50%) lower than expected.



Coldman A, Phillips N, Wilson C, Decker K, Chiarelli AM, Brisson J, Zhang B, Payne J, Doyle G, Ahmad R. Pan-canadian study of mammography screening and mortality from breast cancer. J Natl Cancer Inst. 2014 Oct 1;106(11).

# Observational studies show that the decline in deaths is linked to screening.

1. Tabar L, Vitak B, Tony HH, Yen MF, Duffy SW, Smith RA. Beyond randomized controlled trials: organized mammographic screening substantially reduces breast carcinoma mortality. Cancer 2001;91:1724-31

 Kopans DB. Beyond Randomized, Controlled Trials: Organized Mammographic Screening Substantially Reduces Breast Cancer Mortality. Cancer 2002;94: 580-581
 Duffy SW, Tabar L, Chen H, Holmqvist M, Yen M, Abdsalah S, Epstein B, Frodis Ewa, Ljungberg E, Hedborg-Melander C, Sundbom A, Tholin M, Wiege M, Akerlund A, Wu H, Tung T, Chiu Y, Chiu Chen, Huang C, Smith RA, Rosen M, Stenbeck M, Holmberg L. The Impact of Organized Mammography Service Screening on Breast Carcinoma Mortality in Seven Swedish Counties. Cancer 2002;95:458-469.

4. Otto SJ, Fracheboud J, Looman CWN, Broeders MJM, Boer R, Hendriks JNHCL, Verbeek ALM, de Koning HJ, and the National Evaluation Team for Breast Cancer Screening\* Initiation of population-based mammography screening in Dutch municipalities and effect on breast-cancer mortality: a systematic review Lancet 2003;361:411-417.
5 Swedish Organised Service Screening Evaluation Group. Reduction in breast cancer mortality from organized service screening with mammography: 1. Further confirmation with extended data. Cancer Epidemiol Biomarkers Prev. 2006;15:45-51
6. Coldman A, Phillips N, Warren L, Kan L. Breast cancer mortality afterscreening mammography in British Columbia women. Int J Cancer. 2007 Mar

1;120(5):1076-80.

# Observational studies show that the decline in deaths is linked to screening.

7. Jonsson H, Bordás P, Wallin H, Nyström L, Lenner P. Service screening with mammography in Northern Sweden: effects on breast cancer mortality - an update. J Med Screen. 2007;14(2):87-93.

8. Paap E, Holland R, den Heeten GJ, et al. A remarkable reduction of breast cancer deaths in screened versus unscreened women: a case-referent study. Cancer Causes Control 2010; 21: 1569-1573

 Otto SJ, Fracheboud J, Verbeek ALM, Boer R, Reijerink-Verheij JCIY, Otten JDM, Broeders MJM, de Koning HJ, and for the National Evaluation Team for Breast Cancer Screening. Mammography Screening and Risk of Breast Cancer Death: A Population-Based Case–Control Study. Cancer Epidemiol Biomarkers Prev. Published OnlineFirst December 6, 2011; doi: 10.1158/1055-9965.EPI-11-0476

. 10. van Schoor G, Moss SM, Otten JD, Donders R, Paap E, den Heeten GJ, Holland R, Broeders MJ, Verbeek AL. Increasingly strong reduction in breast cancer mortality due to screening. Br J Cancer. 2011 Feb 22. [Epub ahead of print]

11. Mandelblatt JS, Cronin KA, Bailey S, et.al. Effects of mammography screening under different screening schedules: model estimates of potential benefits and harms. Annals of Internal Medicine, 2009; 151: 738-747; see also http://cisnet.cancer.gov, last accessed 16 April 2011.



# Observational studies show that the decline in deaths is linked to screening.

12.Hellquist BN, Duffy SW, Abdsaleh S, Björneld L, Bordás P, Tabár L, Viták B, Zackrisson S, Nyström L, Jonsson H. Effectiveness of population-based service screening with mammography for women ages 40 to 49 years: evaluation of the Swedish Mammography Screening in Young Women (SCRY) cohort. Cancer. 2011 Feb 15;117(4):714-22

3. Broeders M, Moss S, Nyström L, Njor S, Jonsson H, Paap E, Massat N, Duffy S, Lynge E, Paci E; EUROSCREEN Working Group. The impact of mammographic screening on breast cancer mortality in Europe: a review of observational studies. J Med Screen. 2012;19 Suppl 1:14-25. Review

14. Hofvind S, Ursin G, Tretli S, Sebuødegård S, Møller B. Breast cancer mortality in participants of the Norwegian Breast Cancer Screening Program. Cancer. 2013 Sep 1;119(17):3106-12

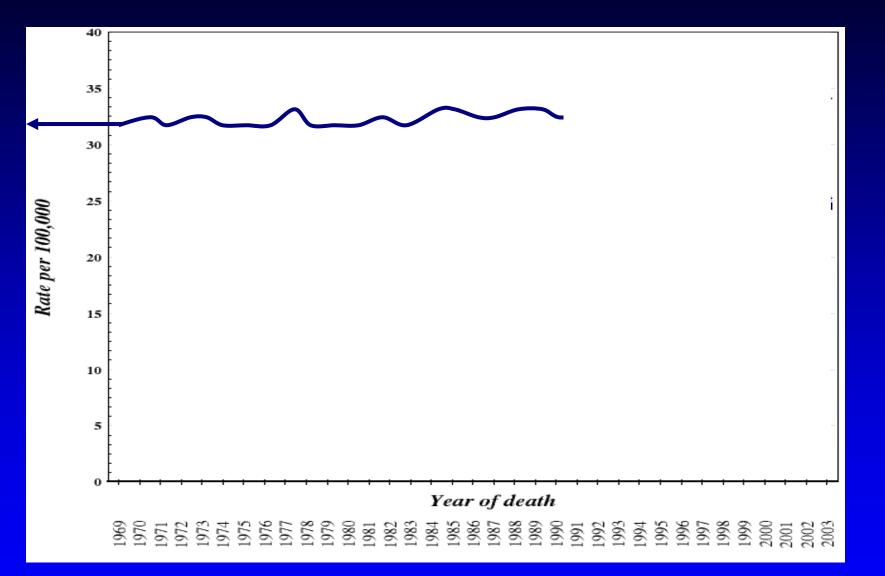
15. Sigurdsson K, Olafsdóttir EJ. Population-based service mammography screening: the Icelandic experience. Breast Cancer (Dove Med Press). 2013 May 9;5:17-25 16. Coldman A, Phillips N, Wilson C, Decker K, Chiarelli AM, Brisson J, Zhang B, Payne J, Doyle G, Ahmad R. Pan-canadian study of mammography screening and mortality from breast cancer. J Natl Cancer Inst. 2014 Oct 1;106(11).



## IN THE U.S.

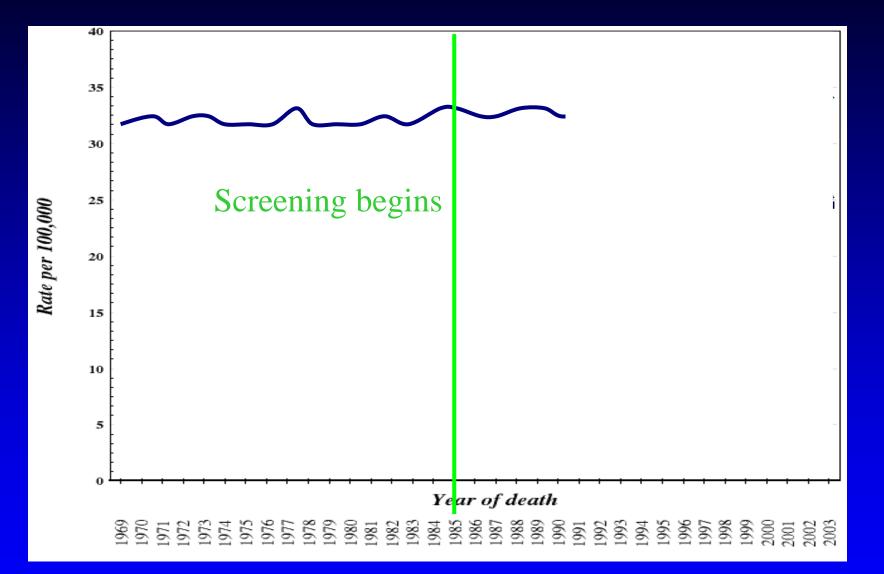


#### Breast Cancer Death Rate 1969-1990



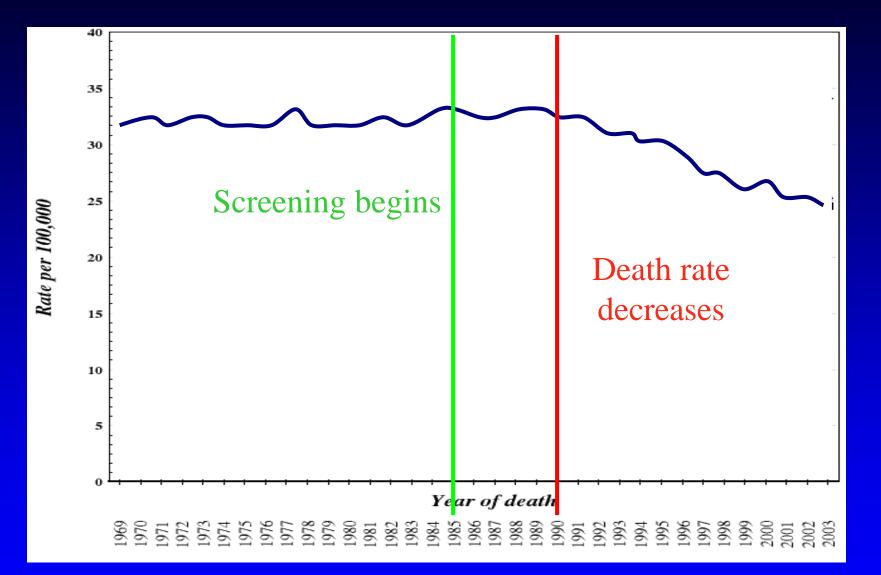
Surveillance, Epidemiology, and End Results (SEER) Program (www.seer.cancer.gov) SEER\*Stat Database: Mortality - All COD, Public-Use With State, Total U.S. (1969-2003), National Cancer Institute, DCCPS, Surveillance Research Program, Cancer Statistics Branch, released April 2006. Underlying mortality data provided by NCHS (www.cdc.gov/nchs).

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#### **Breast Cancer Deaths Averted Over 3 Decades**

R. Edward Hendrick, PhD <sup>(D)</sup>; Jay A. Baker, MD<sup>2</sup>; and Mark A. Helvie, MD<sup>3</sup>

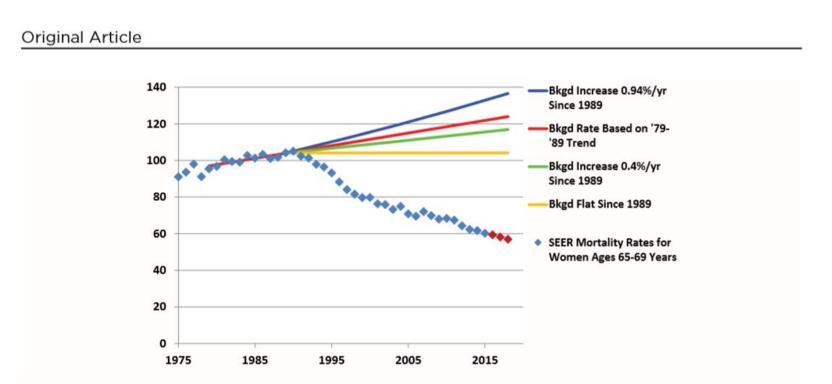
**BACKGROUND:** From 1975 to 1990, female breast cancer mortality rates in the United States increased by 0.4% per year. Since 1990, breast cancer mortality rates have fallen between 1.8% and 3.4% per year, a decrease that is attributed to increased mammography screening and improved treatment. **METHODS:** The authors used age-adjusted female breast cancer mortality rate and population data from the Surveillance, Epidemiology, and End Results (SEER) program to estimate the number of breast cancer deaths averted by screening mammography and improved treatment since 1989. Four different assumptions regarding background mortality rates (in the absence of screening mammography and improved treatment) were used to estimate deaths averted for women aged 40 to 84 years by taking the difference between SEER-reported mortality rates and background mortality rates for each 5-year age group, multiplied by the population for each 5-year age group. SEER data were used to estimate annual and cumulative breast cancer deaths averted in 2012 and 2015 and extrapolated SEER data were used to estimate deaths averted in 2018. **RESULTS:** The number of single-year breast cancer mortality reductions ranged from 38.6% to 50.5% in 2012, from 41.5% to 54.2% in 2015, and from 45.3% to 58.3% in 2018. Cumulative breast cancer deaths averted since 1989 ranged from 237,234 to 370,402 in 2012, from 305,934 to 483,435 in 2015, and from 384,046 to 614,484 in 2018. **CONCLUSIONS:** Since 1989, between 384,000 and 614,500 breast cancer deaths have been averted through the use of mammography screening and improved treatment. **Cancer 2018;0:1-7**. © *2018 American Cancer Society*.

KEYWORDS: breast cancer, female, mortality rate, screening mammography, therapy, treatment, United States.

(Hendrick RE, Baker JA, Helvie MA. Breast cancer deaths averted over 3 decades. Cancer. 2019 Feb 11. doi: 10.1002/cncr.31954. [Epub ahead of print])



#### Between 384,000 and 614,000 lives have been saved since 1990 based on what the projected mortality would have been.



**Figure 3.** Surveillance, Epidemiology, and End Results (SEER) age-adjusted mortality rates (per 100,000 women) for women aged 65 to 69 years by year from 1975 through 2015 (blue data points). Dark red data points represent linear extrapolation of the mortality rate to 2016 through 2018 based on the trend from 2006 through 2015. Solid lines represent the 4 different assumptions regarding background (Bkgd) mortality rate trends since 1989. Blue indicates increasing by 0.94% per year since 1989; red, based on a linear fit to mortality rates from 1979 through 1989; green, increasing by 0.4% per year since 1989; orange, constant since 1989.



More than 40% fewer women now die each year from breast cancer in large part due to screening and earlier detection

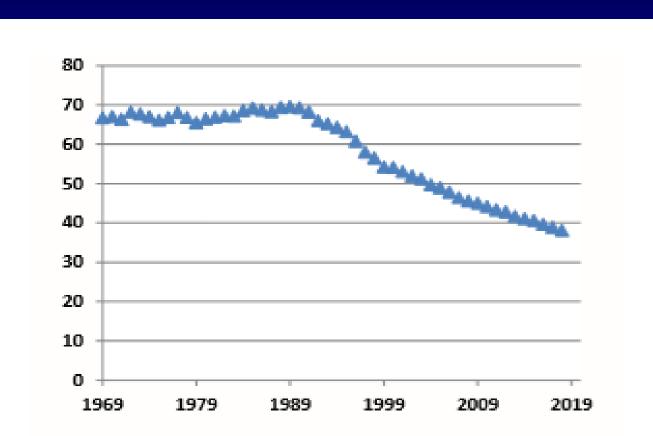


Figure 1. Age-adjusted US breast cancer mortality rates (per 100,000 women) for women aged 40 to 84 years by year from 1969 through 2015 from the Surveillance, Epidemiology, and End Results (SEER) database.



It has been suggested that the decline in breast cancer deaths is due to improvements in therapy.



There certainly have been therapeutic advances, but the numerous observational studies have shown that the death rate declines primarily for women who participate in screening.



**INCIDENCE OF FATAL CANCERS** 

Over 50,000 women ages 40-69:

Prescreening data from 1958-1976 Screening era followed over 39 years 1977-2015

(Tabár L, et al. The incidence of fatal breast cancer measures the increased effectiveness of therapy in women participating in mammography screening. Cancer. 2019 Feb 15;125(4):515-523.])



#### INCIDENCE OF FATAL CANCERS

Despite all women having access to modern therapy, the incidence of mortality was 60% lower at 10 years and 47% lower at 20 years for women participating in screening compared to those who did not.

(Tabár L, et al. The incidence of fatal breast cancer measures the increased effectiveness of therapy in women participating in mammography screening. Cancer. 2019 Feb 15;125(4):515-523.])



#### **ONE FINAL OBSERVATION**

#### The effect of therapy is unclear.

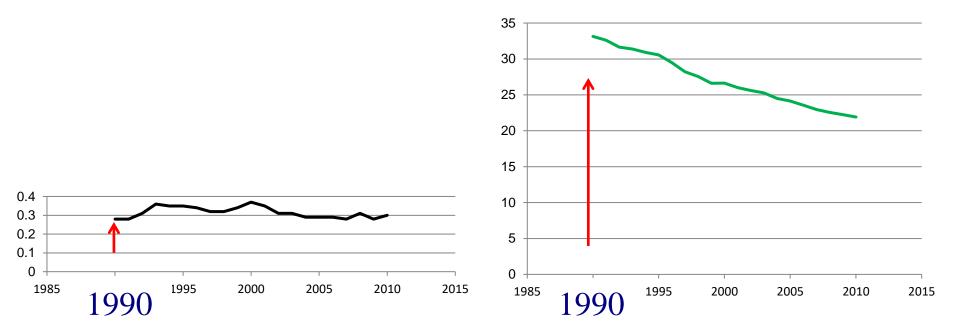
The death rate for men with breast cancer has not declined since 1990 while it has steadily declined for women.



#### UNITED STATES MALES VS FEMALE BREAST CANCER DEATH RATES 1990-2010

#### DEATH RATE PER 100,000 MALES

#### DEATH RATE PER 100,000 FEMALES



Mammography screening began in the mid 1980's and the death rate began to fall in 1990.

Over the same period, with access to the same therapy, the death rate for men increased, then returned to 1990 levels, and has not fallen since 1990.

THE DIFFERENCE ?

#### **WOMEN ARE BEING SCREENED !**



#### **ONE FINAL OBSERVATION**

## What is the difference ?

## Women are being screened and their cancers detected earlier.



#### BREAST CANCER SCREENING

THERE IS ACTUALLY NOW A CONSENSUS BUT IT IS BURIED IN NON-SCIENCE:

All of the major groups agree that the most lives are saved by annual screening starting at the age of 40.



### BREAST CANCER SCREENING

The Panels used results from the National Cancer Institute's 6 computer modeling groups called:

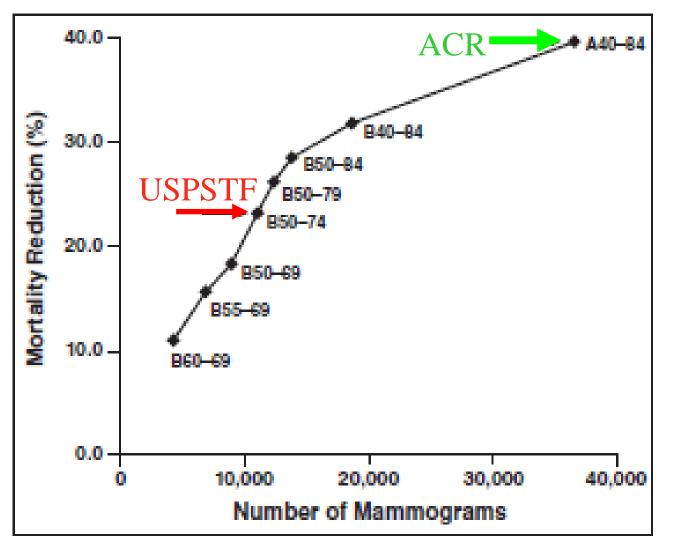
The Cancer Intervention and Surveillance Modeling Network

"CISNET"



#### CISNET SHOWS THAT MOST LIVES ARE SAVED BY ANNUAL SCREENING BEGINNING AT 40

Fig. 1—Percentage mortality reduction from various screening strategies. Note that annual (A) screening from ages 40–84 years (A40–84, *solid arrow*) is estimated to have 71% greater mortality benefit than biennial (B) screening from ages 50–74 years (B50–74, *dashed arrow*). Number of mammograms shown on horizontal axis is per 1,000 women screened. Data shown are mean values of six models from [6].



### The United States Preventive Services Task Force (USPSTF)

"found adequate evidence that mammography screening reduces breast cancer mortality in women ages 40 to 74 years."

(Siu AL; U.S. Preventive Services Task Force. Screening for Breast Cancer: U.S. Preventive Services Task Force Recommendation Statement. Ann Intern Med. 2016 Feb 16;164(4):279-96. doi: 10.7326/M15-2886. Epub 2016 Jan 12. )

## The American College of Physicians (ACP)

"Screening mammography has been shown to decrease the number of deaths from breast cancer in women ages 40-74."

(http://www.acpinternist.org/archives/2012/05/policy.htm)



## The American Cancer Society (ACS)

"Screening mammography in women aged 40 to 69 years is associated with a reduction in breast cancer deaths"

"Women should have the opportunity to begin annual screening between the ages of 40 and 44 years".



#### THE PANELS

Experts in screening were specifically excluded from the panels! None of the panelists provided care for women with breast cancer.



#### THE PANELS

Unfortunately, the inexpert panels went on to impose their own biases by suggesting that women should delay screening until age 45 (ACS) or age 50 (USPSTF and ACP). Their claim was this would reduce "overdiagnosis", "overtreatment", and "false positives".



#### THE PANELS

Experts need not apply!

By not having experts on the panels allowed "advisors" (opponents of screening) to influence the outcomes.



#### **INEXPERT PANELS**

They claim to have weighed the "harms" (primarily recalls from screening) vs. the "benefits" (lives saved), yet neither group explains:

How many fewer recalls are equivalent to having one woman die, unnecessarily, from breast cancer?



Using the NCI/CISNET models it is estimated that waiting until the age of 45 and going to biennial screening at age 55 (ACS) would mean that 38,000 women now in their thirties would die whose lives could be saved by annual screening starting at the age of 40.



#### USPSTF PANEL IGNORED ITS OWN RESULTS

Using the CISNET models it is estimated that waiting until the age of 50 and screening every two years (USPSTF) would mean that 100,000 women now in their thirties would die whose lives could be saved by annual screening starting at the age of 40.

(Hendrick RE, Helvie MA. USPSTF Guidelines on Screening Mammography Recommendations: Science Ignored. Am. J. Roentgenology 2011; 196: W112 - W116.)

Despite treatment, more than 40,000 women still die each year in the U.S. despite therapy.



In a major study at the Harvard teaching hospitals, where women had access to modern therapy, the majority of the women who died from breast cancer, were not participating in screening.

(Webb, et al. A Failure Analysis of Invasive Breast Cancer Most Deaths From Disease Occur in Women Not Regularly Screened Cancer 2013.)

71% of the breast cancer deaths were among the 20% of women who were not participating in screening.

(Webb, et al. A Failure Analysis of Invasive Breast Cancer Most Deaths From Disease Occur in Women Not Regularly Screened Cancer 2013.)



The Harvard study would suggest that many of the women who die in the U.S. each year were not participating in screening.

Why doesn't our National database (SEER) track how cancers are found?



The fundamental scientific evidence shows that the most lives are saved by annual screening starting at the age of 40

## PERIOD



#### BREAST CANCER IS NOT A TRIVIAL PROBLEM FOR WOMEN IN THEIR FORTIES

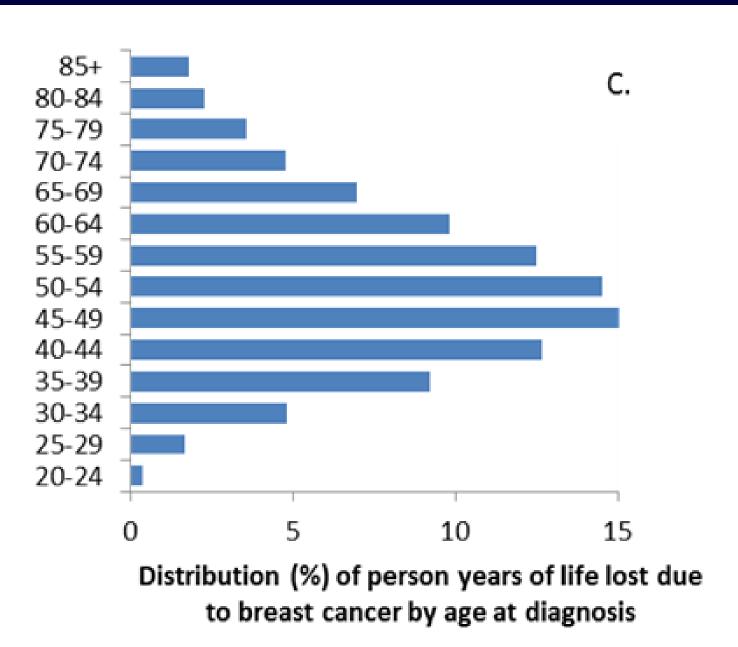


#### BREAST CANCER IS NOT A TRIVIAL PROBLEM FOR WOMEN IN THEIR FORTIES

More than 30,000 women are diagnosed with breast cancer each year while in their forties.



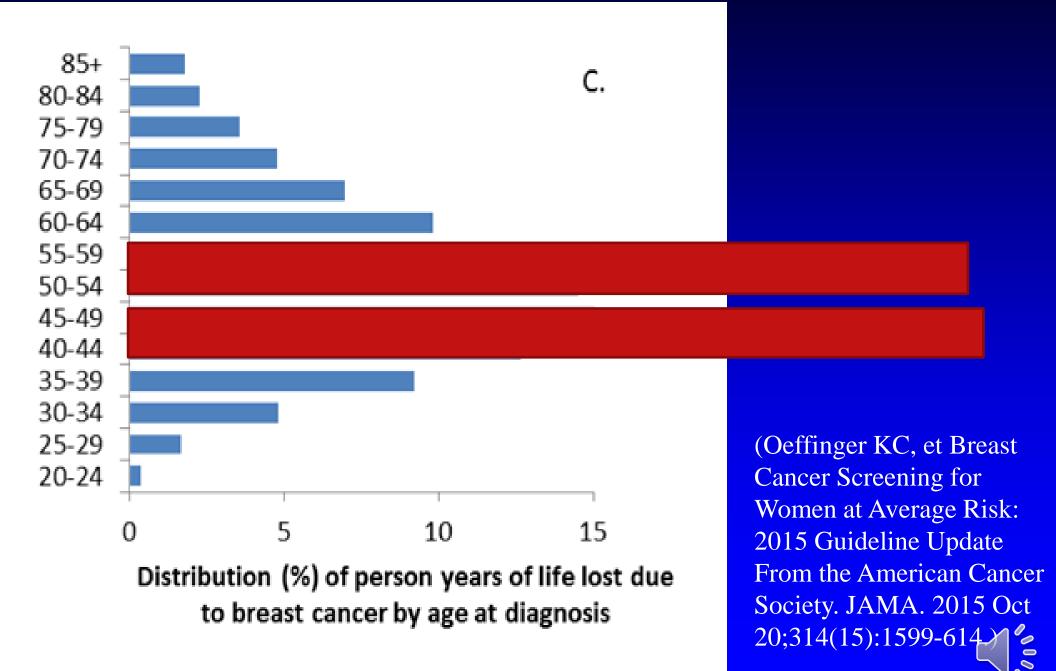
#### Years of Life Lost to Breast Cancer



(Oeffinger KC, et Breast Cancer Screening for Women at Average Risk: 2015 Guideline **Update From** the American Cancer Society. JAMA. 2015 Oct 20;314(15):1599 -614.)



## There are more years of life lost to breast cancer for women ages 40-49 than women ages 50-59



#### BREAST CANCER IS NOT A TRIVIAL PROBLEM FOR WOMEN IN THEIR FORTIES

There are more years of life lost to breast cancer among women ages 40-49 than among women ages 50-59.

Oeffinger KC, et al Breast Cancer Screening for Women at Average Risk: 2015 Guideline Update From the American Cancer Society. JAMA. 2015 Oct 20;314(15):1599-614.



## BREAST CANCER SCREENING WHY THE CONTROVERSIES ?

Since the issues have not changed, and they have all been addressed, scientifically, the continued use of misinformation to deny women access to screening is either due to a failure to understand the data and legitimate scientific analysis, or a malicious effort to mislead.



The effort to reduce access to screening has been going on steadily for decades. We had "ALTERNATIVE FACTS" before they became mainstream!



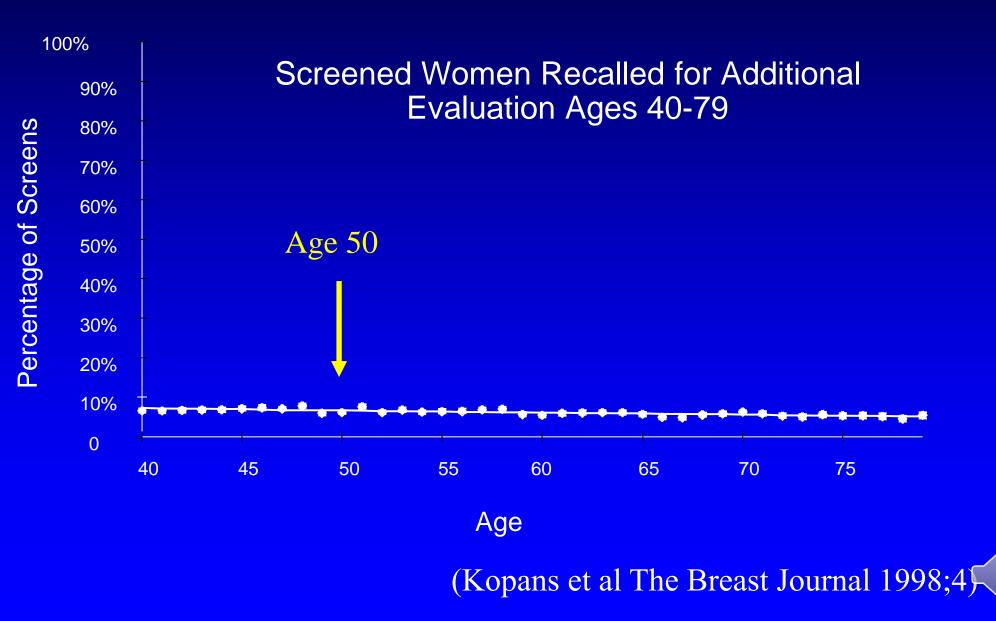
BREAST CANCER SCREENING "ALTERNATIVE FACTS"

There are NO DATA (ZERO) to support the use of the age of 50 as a threshold for screening.

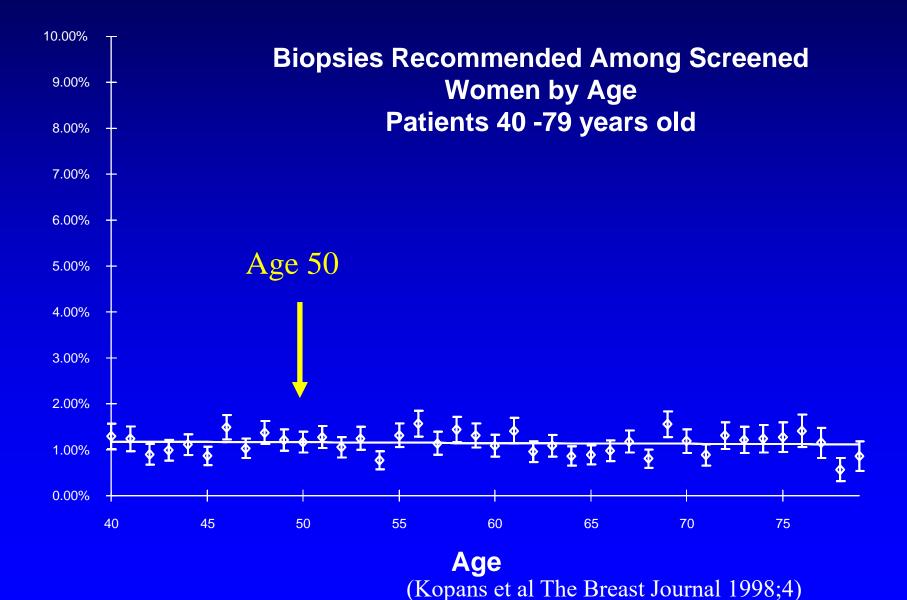
None of the parameters of screening change abruptly at the age of 50 or any other age.



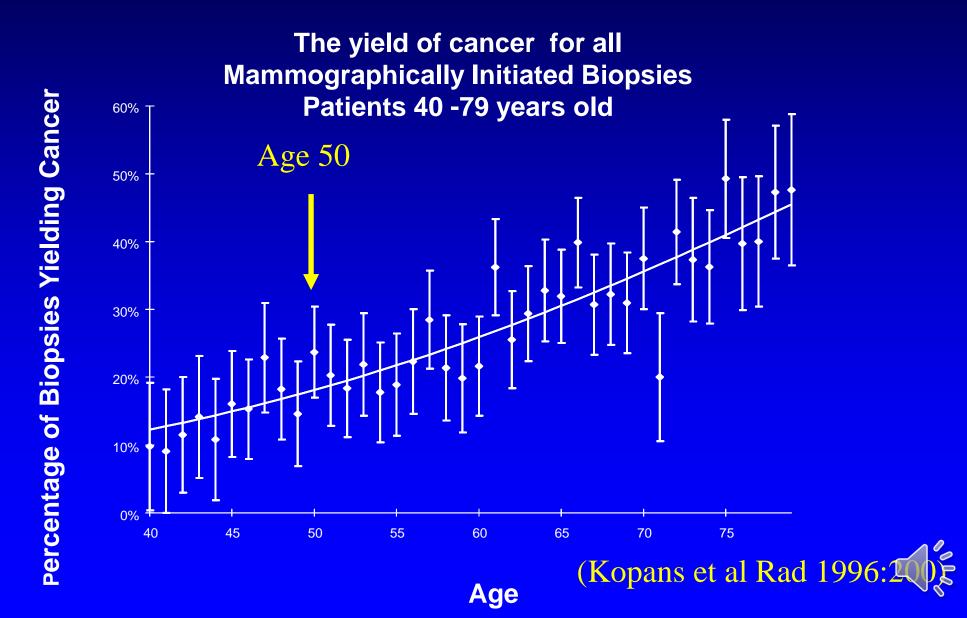
# The recall rate from screening decreases gradually with increasing age from 8% to 6% with no abrupt change at age 50 or any other age



The percentage of women who are recommended for biopsy is fairly constant with no abrupt change at age 50 or any other age.

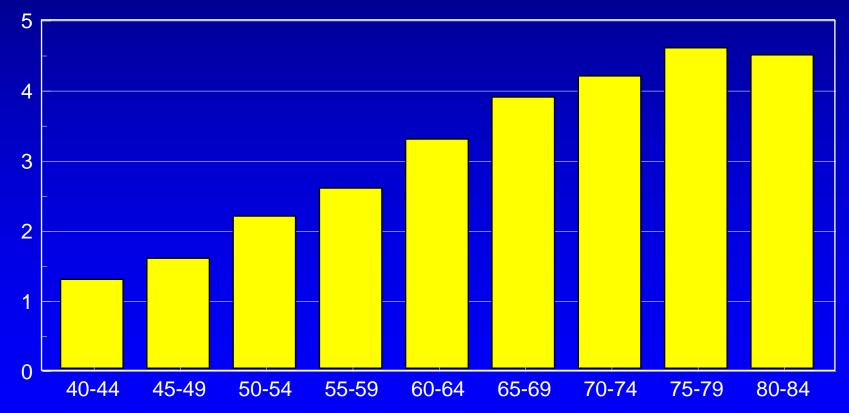


The positive predictive value of a biopsy instigated by mammography goes up with the prior probability of cancer in the population with no abrupt change at any age.



#### ANNUAL BREAST CANCER INCIDENCE (per 1000) BY AGE

CANCERS/1000/YEAR





#### AGE 50 AND MAMMOGRAPHY SCREENING

#### Fact:

The cancer detection rate increases steadily with increasing age along with the steady increase in breast cancer incidence, reflecting the prior probability of breast cancer that increases with age.

<u>There is no abrupt change at age 50 or</u> any other age.

### BREAST CANCER SCREENING

How to make it appear as if the cancer detection rate changes suddenly at the age of 50.



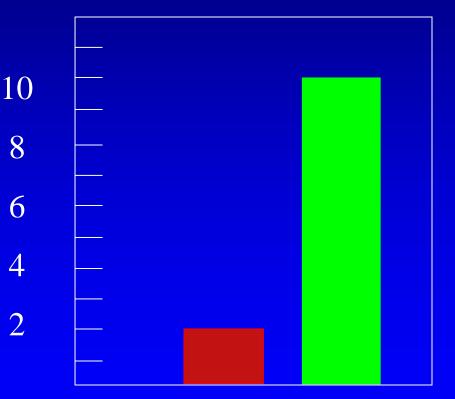
#### **A SIMPLE WAY TO BIAS CONCLUSIONS**

#### (Kerlikowske et al – UCSF-JAMA 1993)

Compared women ages 30-49 to all women ages 50-70+

Cancers per 1000 women

2



30-49 50-70+



#### A SIMPLE WAY TO BIAS CONCLUSIONS

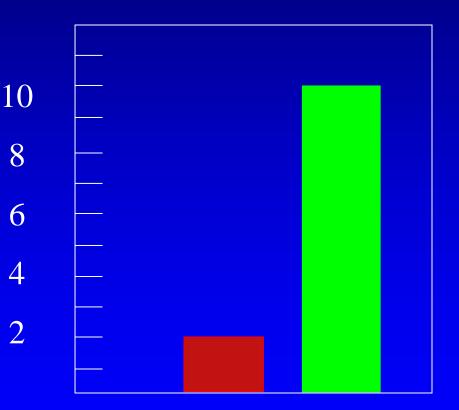
#### (Kerlikowske et al – UCSF-JAMA 1993)

Compared women ages 30-49 to all women ages 50-70+

Cancers per 1000 women

2

4



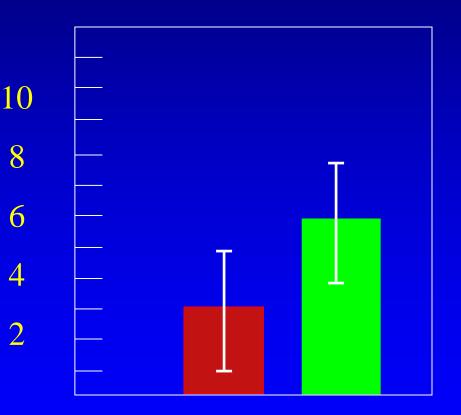
30-49



50-70+

## USING A MORE APPROPRIATE COMPARISON (40-49 VS. 50-59) THERE IS LITTLE DIFFERENCE

3 per 1000 for women ages 40-49 and 6 per 1000 for women ages Cancer 50-59. s per 1000 With overlapping women confidence intervals there is no significant difference

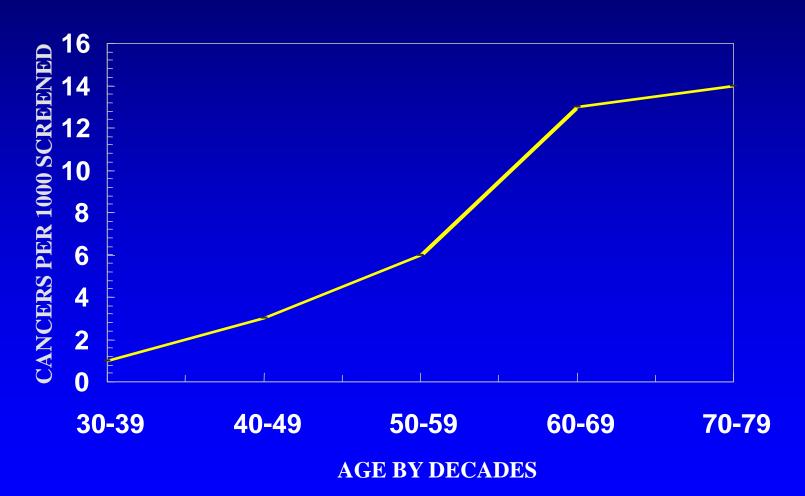


40-49 50-59



### MORE APPROPRIATE AGE GROUPING BY DECADE

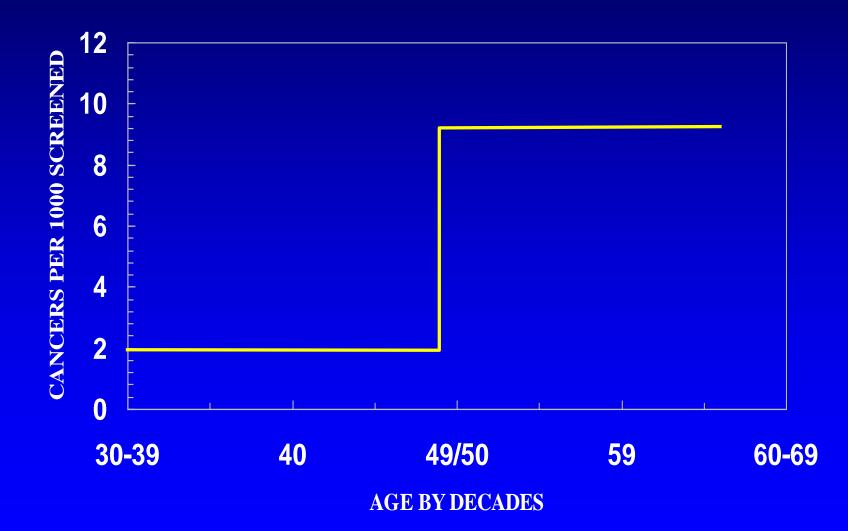
Kerlikowske et al - JAMA 1993





## BIASING DATA BY INAPPROPRIATE AGE GROUPING

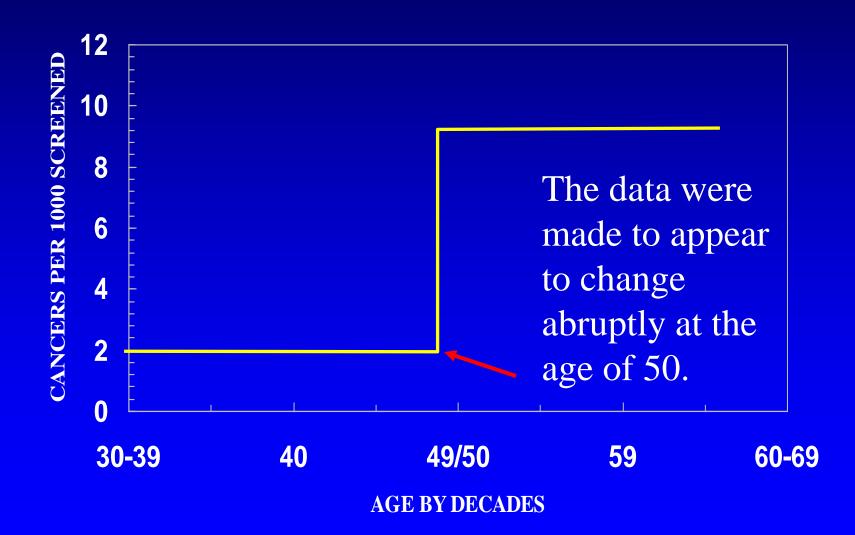
Kerlikowske et al - JAMA 1993





#### BIASING DATA BY INAPPROPRIATE AGE GROUPING

Kerlikowske et al - JAMA 1993





### INAPPROPRIATE DATA GROUPING CAN BE GREATLY MISLEADING

"The yield [of cancers] of the first mammogram was five times higher in women 50 years of age and older (10 cancers per 1000 studies compared with 2 cancers per 1000 studies)... Clearly mammography is much more efficient in detecting breast cancers in older women."

(Sox - Annals of Int Med:1995)



# HOW YOU WERE MISLED

FACT: Opponents of screening women ages 40-49 have repeatedly grouped them together as if they are a uniform group and compared them to the group of <u>all</u> women ages 50 and over as if they are a uniform group. This also takes factors that change gradually with increasing age and makes them appear to change suddenly at the age of 50.

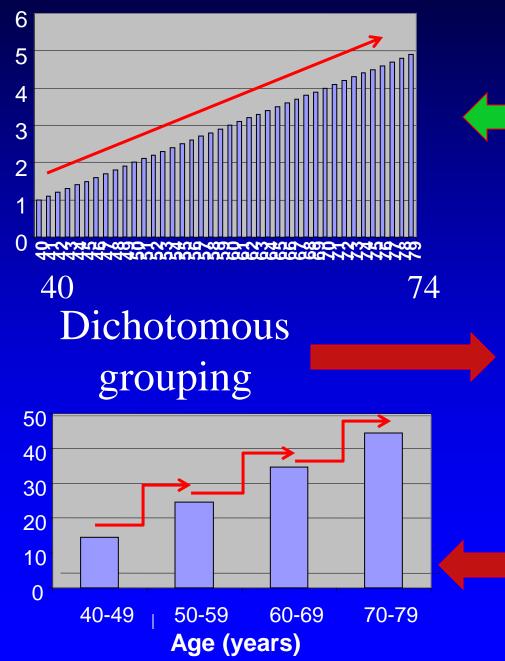


## WHY IS THE MYTH PERPETUATED ??

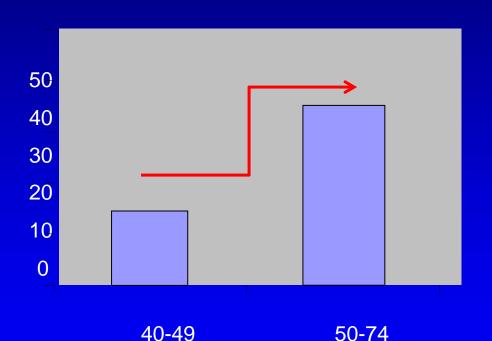
The age of 50 has been imbued with importance by scientifically unjustified subgroup analyses, and dichotomous data grouping that makes steady changes appear to change at the age of 50. Investigators should know better.



Age grouping has been used to make data that actually change gradually with increasing age appear to change suddenly at the age of 50.



Reality = continuous gradual change



USPTSF = group by decade

#### AGE 50 AND MAMMOGRAPHY SCREENING

The suggestion that any of the parameters of screening change abruptly at the age of 50 is a myth that is unsupported by any science. Women should be informed, and investigators should cease grouping data to make age 50 appear as if it has any true importance.



# MAMMOGRAPHY SCREENING

# **RADIATION RISK**



# MAMMOGRAPHY SCREENING

One of the recurring issues that has been used to dissuade women from mammography screening is the concern over risk of carcinogenesis from radiation.



# The Origin of The Breast Cancer Radiation Fear

In 1976, Bailar suggested that radiation from mammograms would cause more cancers deaths than would be cured.

(Bailar, JC. Ann Intern Med 1976 84:77-84

Bailar was off by at least an order of magnitude.

The concerns did lead to valuable improvements in technology and a marked reduction in mammographic doses, but exaggerated concerns have persisted.



Misinformation has been perpetuated in journals that are monitored by the media. Susan Love, a "media expert" wrote in the Journal of The American Medical Association, that for every 20,000 mammograms the radiation would cause one breast cancer – just a statement – no reference.

(Lee Davis D, Love SM. Mammographic Screening. JAMA 1994;271:152-153.)



Any exaggerated estimate of radiation risk from mammography was reinforced by Berry in his summary of the Consensus Development Conference in 1997 on mammography screening for women in their forties. This was passed to the public by the media.

> (Kolata G. Stand on Mammograms Greeted by Outrage. New York Times. Tuesday January 28, 1997.)

# RADIATION RISK TO THE BREAST IS TRIVIAL COMPARED TO THE BENEFIT

The major problem with mammography screening is not the radiation risk, which is unmeasurable and small, but, rather, the repeated dissemination of misinformation.



# MISINFORMATION

"Large studies have found that earlier mammograms save almost no lives; since the radiation can cause cancer it therefore makes sense to minimize them"

Sharon Begley – Newsweek Dec. 10, 2009



# RADIATION RISK TO THE BREAST

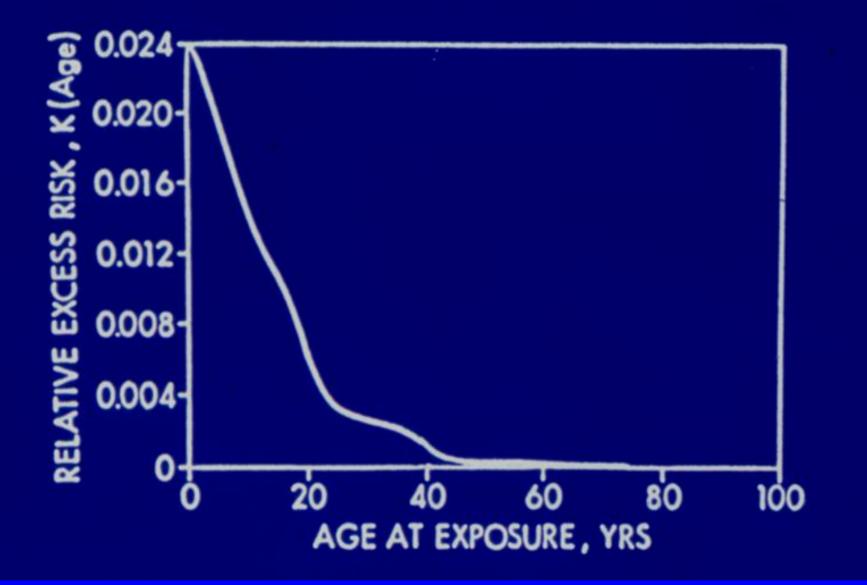
# Fact:

Radiation risk to the breast is related to the age at exposure, and drops off rapidly with increasing age.

(NCRP Report No. 149;2004)



#### RADIATION RISK DECREASES WITH INCREASING AGE





BREAST CANCERS MAY BE INTIATED EARLY IN LIFE

## Fact:

Women treated for Hodgkins disease, with mantle radiation, while in their teens, have a 35% chance of developing breast cancer. The same treatment after the age of 30 results in no excess breast cancer.

(Bhatia et al. New Engl J Med 1996;334:745-751.)

# RADIATION RISK TO THE BREAST IS AGE RELATED

The undifferentiated breast, in teenage women (and younger), is susceptible to radiation carcinogenesis.

There are no direct data (it is all extrapolated) to show any risk from mammograms for women ages 40 and over.

# BREAST CANCERS MAY BE INTIATED EARLY IN LIFE

The developed breast (cellular differentiation) appears to be much less susceptible to radiation carcinogenesis.

? Concentration of stem cells ?



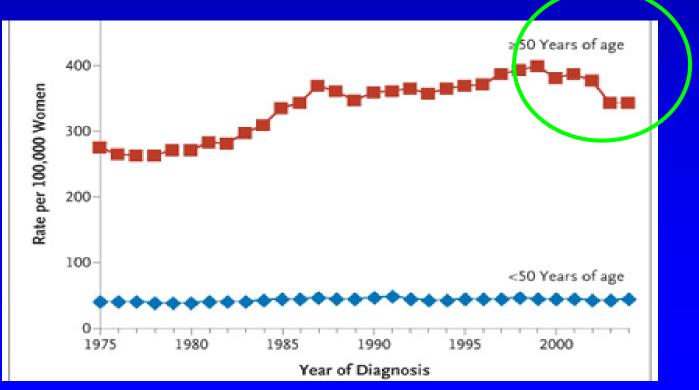
# RADIATION RISK TO THE BREAST IS AGE RELATED

The theoretical risk from mammograms, extrapolated from high dose data, is outweighed by even a small benefit.

(Mettler FA, Upton AC, Kelsey CA, Rosenberg RD, Linver MN. Benefits versus Risks from Mammography: A Critical Assessment. Cancer 1996;77:903-909.)



Hundreds of millions of mammograms have been performed since 1980. If mammograms were causing cancer, the incidence would be increasing among women in their fifties. The incidence of breast cancer has decreased in these women.



# THE "HARMS" OF SCREENING A pejorative term introduced by those interested in reducing access to screening.



## THE "HARMS" OF SCREENING

# "FALSE POSITIVES"

The effort to reduce access to screening has emphasized the "harms" of screening. These are dominated by the "false positives".

What they fail to explain is that most "false positives" are women who are recalled from screening for additional evaluation and do not have cancer. THE "HARMS" OF SCREENING RECALLS FROM SCREENING 1000 women screened

100 (10%) recalled for additional evaluation (which is the same as for Pap testing) 65 have a few images or an ultrasound and nothing is found. 26 are asked to return in 6 months 19 have an image guided needle biopsy using local anesthesia 5-8 have breast cancer



## THE "HARMS" OF SCREENING

19 women out of 1000 have an imaging guided needle biopsy under local anesthesia and 20-40% are found to have breast cancer.

Before there was imaging and surgeons biopsied areas of clinical concern, the yield of cancers was lower (15%) and palpable cancers are larger and later stage, and less likely to be cured than those detected by mammography.

(Spivey GH, Perry BW, Clark VA, & et al, Predicting the Risk of Cancer at the Time of Breast Biopsy. The American Surgeon 1982;48 No.7: 326-332)



WE HAVE A MAJOR PROBLEM !

The major medical journals are preventing an open discussion of important medical issues, and the media have been taught that we are a biased vested interest



#### UNETHICAL BEHAVIOR BY MAJOR MEDICAL JOURNALS

Journals such as:

The New England Journal of Medicine The Annals of Internal Medicine The Journal of the American Medical Association "The Journal of the National Cancer Institute" (which is not the NCI's journal)

Have refused to publish work supporting screening while publishing papers opposing screening (particularly for women ages 40-49)

## THE "HARMS" OF SCREENING

The argument to reduce access to screening now revolves around the "Harms" of screening.

"False Positives"
 "Overdiagnosis"
 "Overtreatment"



"OVERDIAGNOSIS" AND "OVERTREATMENT" It is a diversion to blame these on screening

Pathologists determine the diagnosis. It is up to them to correctly identify potentially lethal cancers.

Treatment is decided by oncologists who know that they have to overtreat many women because they cannot accurately predict who will benefit.

Blaming screening for "overdiagnosis" and "overtreatment" is like blaming the engines in our cars for traffic accidents!

## THE "HARMS" OF SCREENING

The only "harm" that can be reduced by delaying screening are the "false positives" and this is a misnomer. Women are not being told that they have cancer when they do not. These are simply "recalls" from screening when women are reassured that everything is fine. They are, pejoratively, called "false positives"!

## "OVERDIAGNOSIS "

The claim is that screening finds cancers that would never become clinically significant. The panels that seek to limit screening argue that these cancers would go away on their own so that waiting until the age of 50 and screening every two years will reduce "overdiagnosis" and hence the "overtreatment" of these fake cancers.

## "FAKE BREAST CANCERS"

Those seeking to limit access to screening argue that there are "fake" cancers that would go away if left undetected. This is beyond nonsense.

There have been a handful of cancers that have gone away on their own ("miracles" certainly not a common event). Some of the women still died from their "disappearing" cancers!



## "FAKE BREAST CANCERS"

The hypocrisy is that the effort is to reduce access to screening despite the fact that the <u>miraculously disappearing</u> <u>cancers were all clinically evident!</u>

Perhaps we should stop treating clinically evident cancers because they may disappear on their own!



## PROMULGATING MISINFORMATION

## "FAKE BREAST CANCERS"

There is not a single credible report of an invasive breast cancer, detected by mammography "melting away" on its own. If this occurred as frequently as 30-50% of the time someone should have at least seen a few cases!!



#### THE "NORDIC COCHRANE CENTER" HAS PROMULGATED MISINFORMATION

#### **OVERDIAGNOSIS**

If there is any 'overdiagnosis" from screening it can only be measured from the randomized, controlled trials, and they have suggested that it is, at most, under 10%, and more likely less than 1%.

1. Zackrisson S, Andersson I, Janzon L, Manjer J, Garne JP. Rate of over-diagnosis of breast cancer 15 years after end of Malmo mammographic screening trial: follow-up study. BMJ. 2006;332:689-92.

 Paci E, Warwick J, Falini P, Duffy SW. Overdiagnosis in screening: is the increase in breast cancer incidence rates a cause for concern? J Med Screen. 2004;11:23-7
 Puliti D, Duffy SW, Miccinesi G, de Koning H, Lynge E, Zappa M, Paci E; EUROSCREEN Working Group. Overdiagnosis in mammographic screening for breast

cancer in Europe: a literature review. J Med Screen. 2012;19 Suppl 1:42-56.



The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

Effect of Three Decades of Screening Mammography on Breast-Cancer Incidence

Archie Bleyer, M.D., and H. Gilbert Welch, M.D., M.P.H.

N Engl J Med 2012;367:1999-2005

Claimed that due to screening in 2008 alone: "breast cancer was overdiagnosed in more than 70,000 women; this accounted for 31% of all breast cancers diagnosed"



# **BIAS IN THE MEDIA**

The next day the New York Times, which has a long history of bias against mammography screening, published an Op Ed piece by Dr. Welch with no rebuttal.

The New York Times

The Opinion Pages

**OP-ED CONTRIBUTOR** 

Cancer Survivor or Victim of Overdiagnosis?

By H. GILBERT WELCH Published: November 21, 2012

Hanover, N.H.



The NEJM paper had no scientific merit and should not have been published.

- 1. They did not have direct patient information but relied on registry summaries.
- 2. They faulted mammography even though they had no idea which women had mammograms and which women had their cancers detected by mammography.
- They, inappropriately, combined DCIS and small invasive cancers calling them "early breast cancer" to dilute the results for invasive cancers

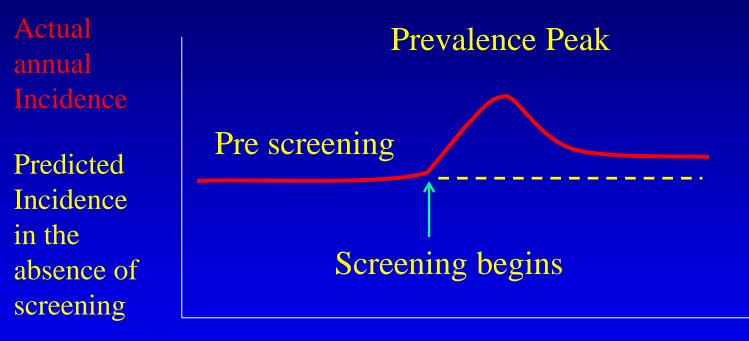


In addition to not having direct patient data, the paper was based on assumptions, estimates, extrapolations, and "guesses" which were simply incorrect.



# SCREENING FUNDAMENTALS

#### Expected Change in Incidence with Screening

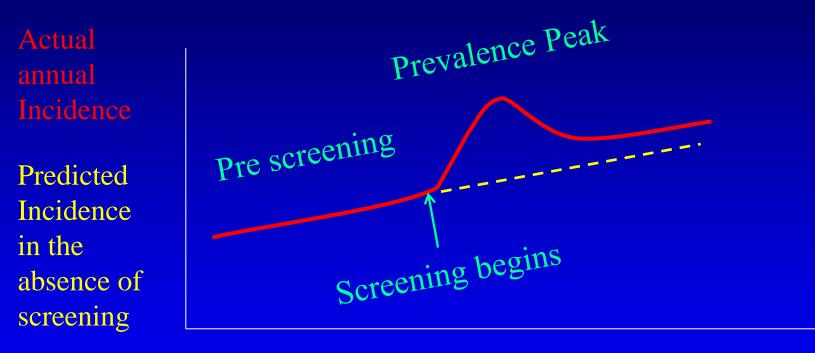


#### Year

With stable screening annual incidence returns to slightly above prescreening levels (leadtime and new prevalence cancers) with cancers at a smaller size

# SCREENING FUNDAMENTALS

#### Expected Change in Incidence with Screening

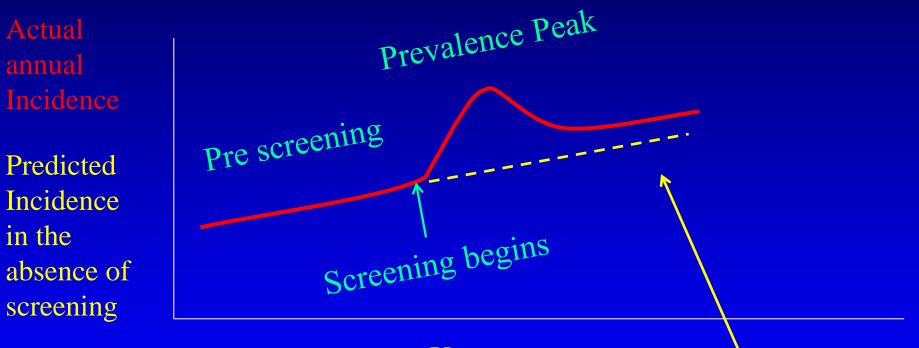


#### Year

What some analysts have misunderstood is that the "baseline" prescreening incidence has been increasing and continued to increase during the screening era  $\sqrt{\frac{2}{2}}$ 

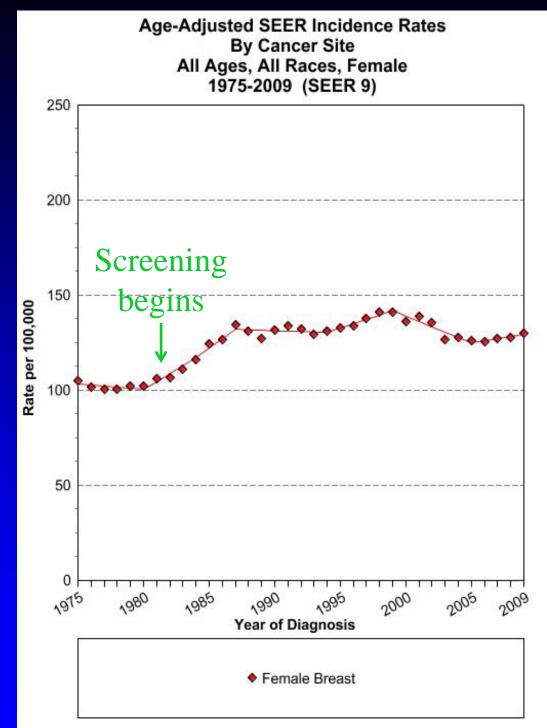
# SCREENING FUNDAMENTALS

#### Expected Change in Incidence with Screening



#### Year

Once screening participation is stable, the incidence will return to an <u>increasing</u> "baseline" if the prescreening incidence had been increasing.



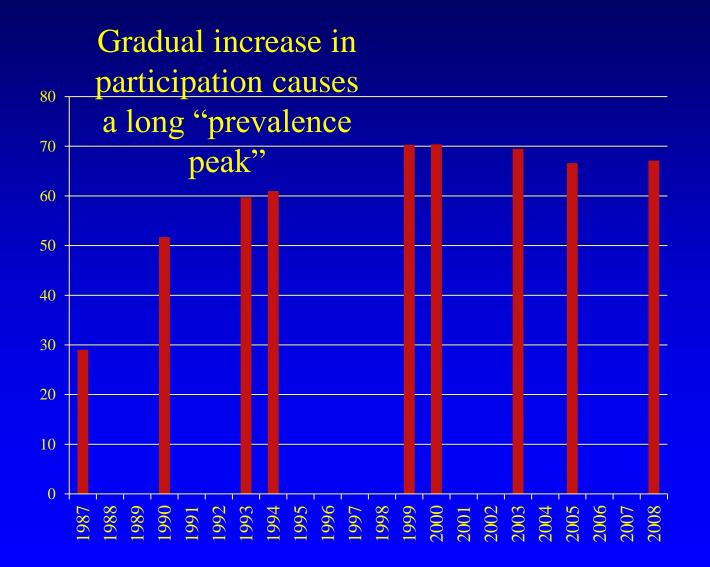
Cancer sites include invasive cases only unless otherwise noted.

Rates are per 100,000 and are age-adjusted to the 2000 US Std Population (19 age groups - Census P25-1130). Regression lines are calculated using the Joinpoint Regression Program Version 3.5, April 2011, National Cancer Institute.

Incidence source: SEER 9 areas (San Francisco, Connecticut, Detroit, Hawaii, Iowa, New Mexico,

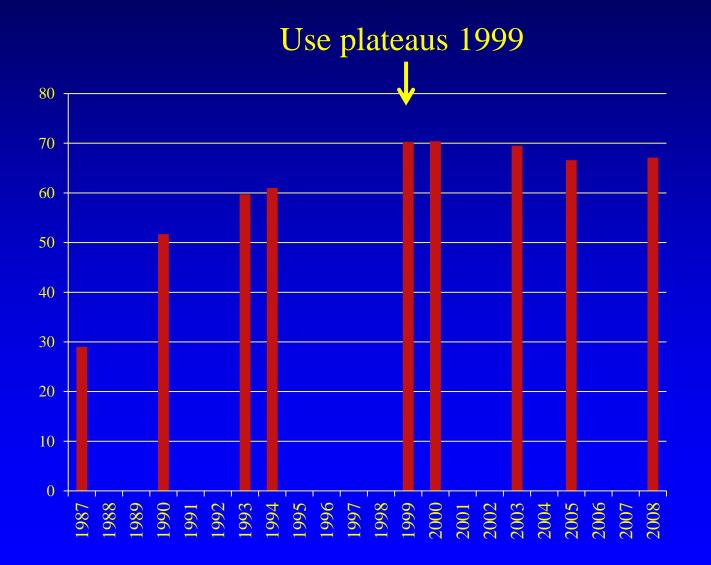


## Use of mammography in the U.S. among women 40 years of age and over http://www.cdc.gov/nchs/data/hus/2010/086.pdf

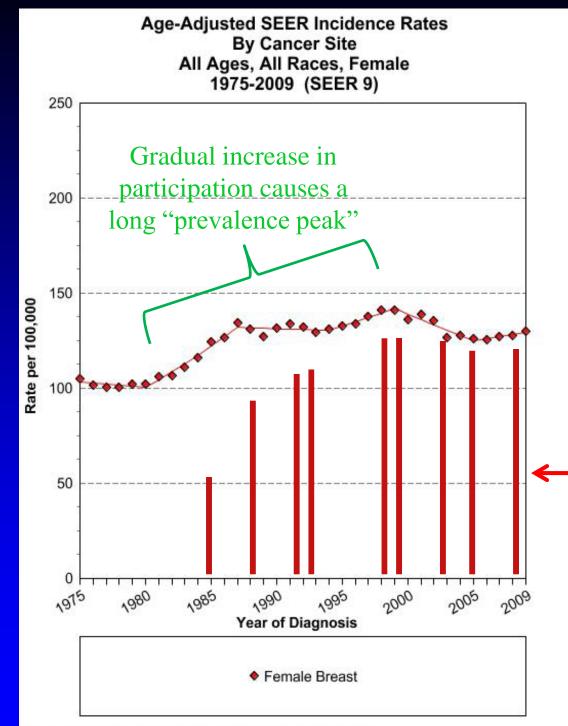




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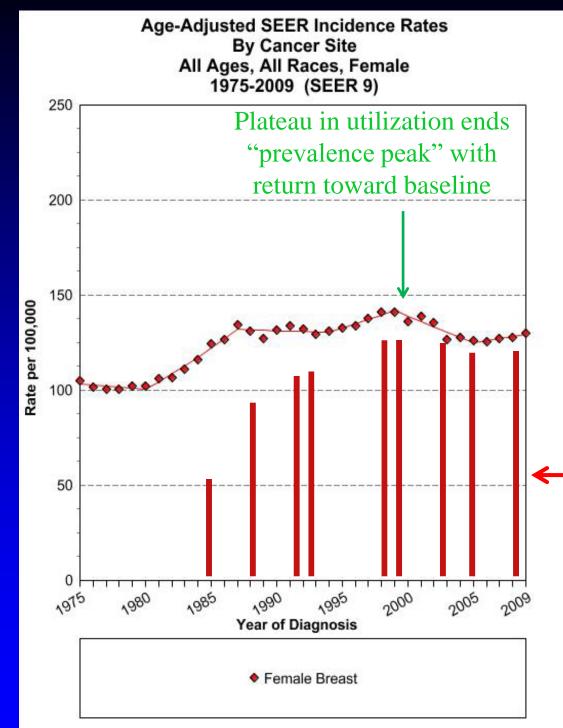
# Utilization of mammography

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# Utilization of mammography

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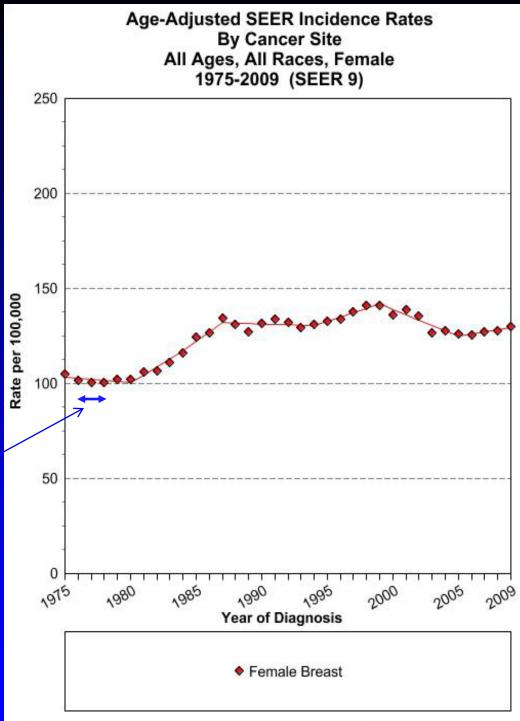
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SEER began in 1973.

**Bleyer and Welch** used data from 1976-1978 to estimate what the baseline breast cancer incidence would have been had screening not been initiated



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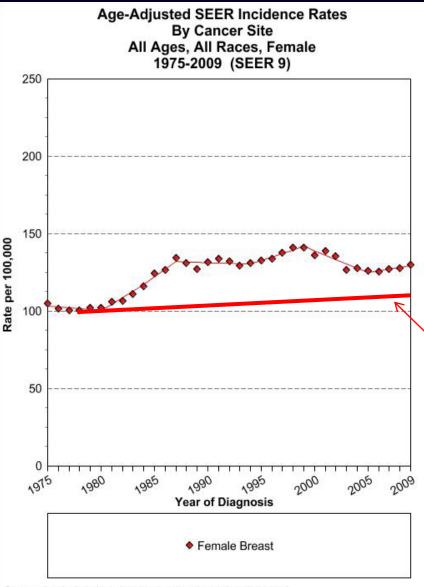
Bleyer and Welch used data from 1976-1978 to estimate what the incidence of breast cancer would have been in 2008 had screening not been initiated in the 1980's.

They ignored the fact that many women were screened after Happy Rockefeller and Betty Ford had breast cancers diagnosed in 1974 and then screening stopped. This was the most unreliable period in the SEER database.

They also ignored 40 years of data.



SEER began in 1973. Bleyer and Welch used data from '76-'78 to estimate that the baseline breast cancer incidence would have increased by 0.25% per year if screening had not been initiated



Cancer sites include invasive cases only unless otherwise noted.

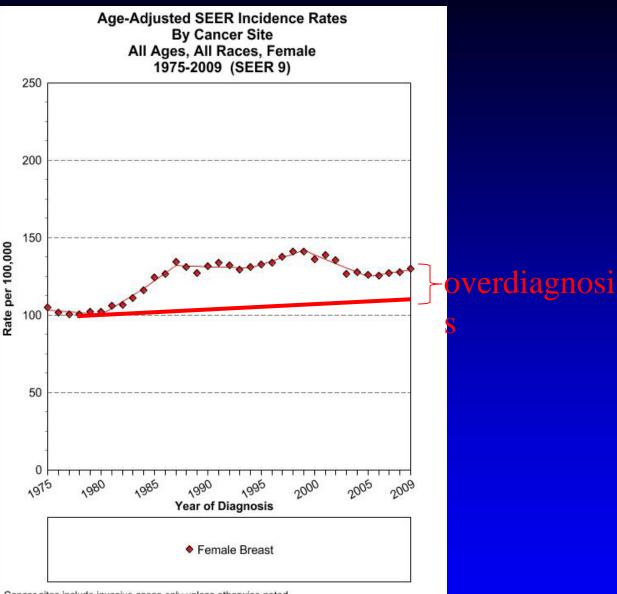
Rates are per 100,000 and are age-adjusted to the 2000 US Std Population (19 age groups -Census P25-1130). Regression lines are calculated using the Joinpoint Regression Program Version 3.5, April 2011, National Cancer Institute.

Incidence source: SEER 9 areas (San Francisco, Connecticut, Detroit, Hawaii, Iowa, New Mexico, Seattle, Utah, and Atlanta).

Bleyer and Welch estimate 0.25% per year baseline increase



**Bleyer and Welch** claim that, since there were more cancers diagnosed in 2008 then they estimated should have occurred in the absence of screening, the excess must be "fake" cancers that would have never been clinically evident.



Cancer sites include invasive cases only unless otherwise noted.

Rates are per 100,000 and are age-adjusted to the 2000 US Std Population (19 age groups -Census P25-1130). Regression lines are calculated using the Joinpoint Regression Program Version 3.5, April 2011, National Cancer Institute.

Incidence source: SEER 9 areas (San Francisco, Connecticut, Detroit, Hawaii, Iowa, New Mexico, Seattle, Utah, and Atlanta).



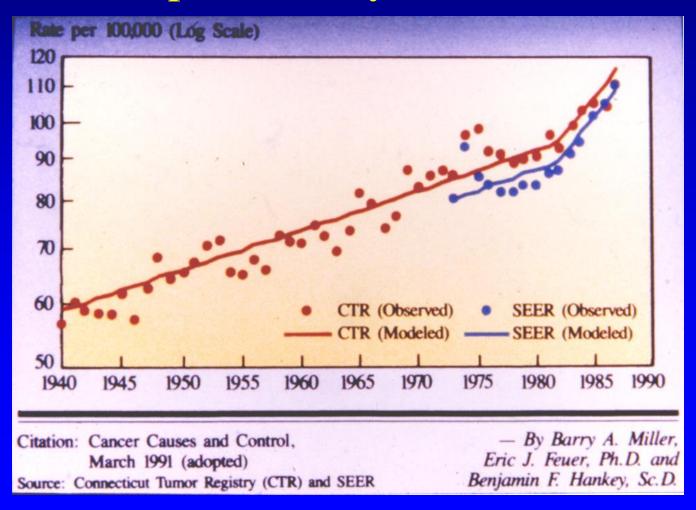
Bleyer and Welch failed to acknowledge that the incidence of invasive breast cancer had been increasing steadily not by 0.25%, but by 1.3% per year since at least 1940.

(Anderson WF, Jatoi I, Devesa SS. Assessing the impact of screening mammography: Breast cancer incidence and mortality rates in Connecticut (1943-2002). Breast Cancer Res Treat. 2006 Oct;99(3):333-40.).



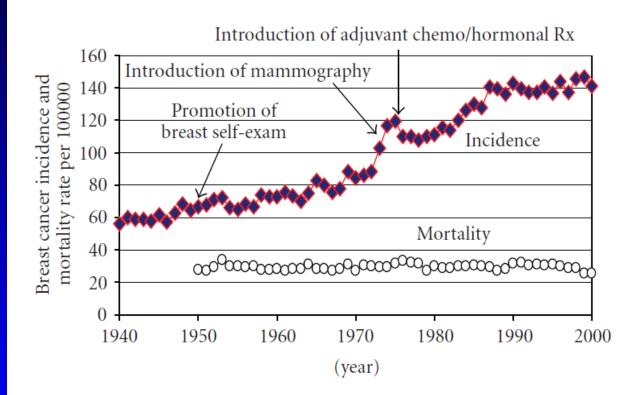
### THE LATEST MISINFORMATION FROM THE DARTMOUTH INSTITUTE ON HEALTH POLICY

In fact, the incidence of invasive breast cancer had been increasing by 1-1.3% each year from 1940 to 1980 prior to any national screening.





## THE INCIDENCE OF BREAST CANCER HAD BEEN INCREASING FOR DECADES



Based on the Connecticut SEER database

FIGURE 1: Breast cancer incidence and mortality, USA, 1940–2000.

(Shulman LN, Willett W, Sievers A, Knaul FM. Breast cancer in developing countries: opportunities for improved survival. J Oncol. 2010;2010)



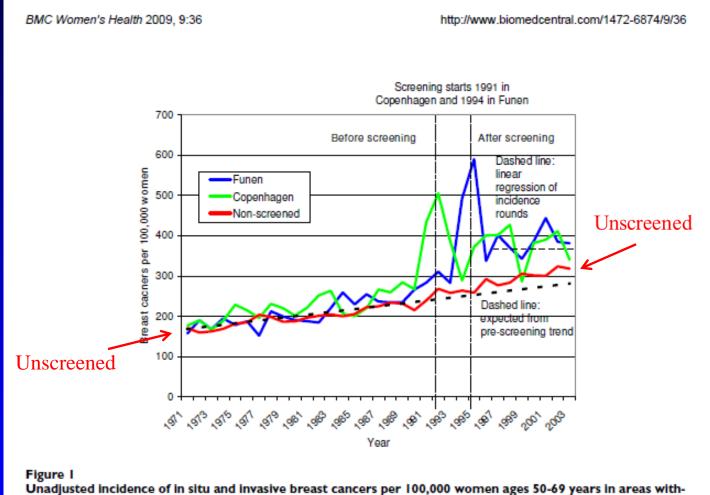
## BLEYER AND WELCH "GUESSED" INCORRECTLY

Every published estimate of the incidence of breast cancer prior to the start of the SEER registry has relied on the data from the Connecticut Tumor Registry.

Shulman LN, Willett W, Sievers A, Knaul FM. Breast cancer in developing countries: opportunities for improved survival. J Oncol. 2010;2010

- Kessler LG, Feuer EJ, Brown ML. Projections of the breast cancer burden to U.S. women: 1990-2000. Prev Med. 1991 Jan;20(1):170-82.
- Garfinkel L, Boring CC, Heath CW Jr. Changing trends. An overview of breast cancer incidence and mortality. Cancer. 1994 Jul 1;74(1Suppl):222-7.
- Miller BA, Feuer EJ, Hankey BF. Recent incidence trends for breast cancer in women and the relevance of early detection: an update. CA Cancer J Clin. 1993 Jan-Feb;43(1):27-41

#### STEADILY INCREASING BACKGROUND INCIDENCE OF UNSCREENED WOMEN IN DENMARK



Onadjusted incidence of in situ and invasive breast cancers per 100,000 women ages 50-69 years in areas out mammography screening and in Copenhagen and in Funen.

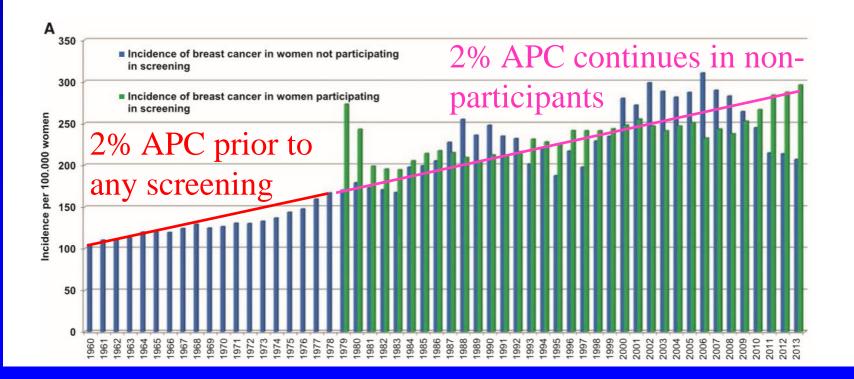
(Jørgensen KJ, Zahl PH, Gøtzsche PC. Overdiagnosis in organised mammography screening in Denmark. A comparative study. BMC Womens Health. 2009 Dec 22;9:36 doi: 10.1186/1472-6874-9-36.)

#### STEADILY INCREASING BACKGROUND INCIDENCE OF UNSCREENED WOMEN IN SWEDEN

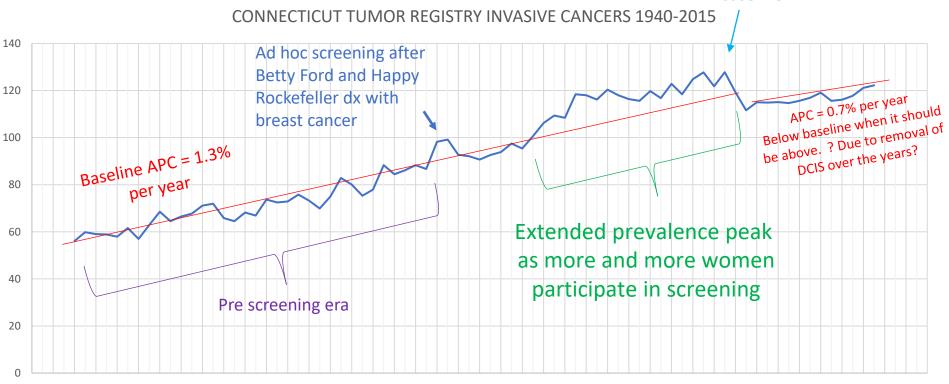
#### The Incidence of Fatal Breast Cancer Measures the Increased Effectiveness of Therapy in Women Participating in Mammography Screening

László Tabár, MD<sup>1</sup>; Peter B. Dean, MD<sup>2</sup>; Tony Hsiu-Hsi Chen, PhD<sup>3</sup>; Amy Ming-Fang Yen, PhD<sup>4</sup>; Sam Li-Sheng Chen, PhD<sup>4</sup>; Jean Ching-Yuan Fann, PhD<sup>5</sup>; Sherry Yueh-Hsia Chiu, PhD<sup>6</sup>; May Mei-Sheng Ku, MSc<sup>3</sup>; Wendy Yi-Ying Wu, PhD<sup>7</sup>; Chen-Yang Hsu, PhD<sup>3</sup>; Yu-Ching Chen, MD<sup>8</sup>; Kerri Beckmann, PhD<sup>9</sup>; Robert A. Smith, PhD<sup>10</sup>; and Stephen W. Duffy, MSc<sup>11</sup>

The Incidence of Fatal Breast Cancer/Tabár et al



# The BASELINE incidence of breast cancer has been increasing since 1940



**Participation plateaus** 

 return toward baseline

Data courtesy of :

1940-1972: Connecticut Tumor Registry

1973-2015: Surveillance, Epidemiology, and End Results (SEER) Program (www.seer.cancer.gov) SEER\*Stat Database: Incidence - SEER 9 Regs Research Data, Nov

# "OVERDIAGNOSIS" IS GROSSLY OVERESTIMATED

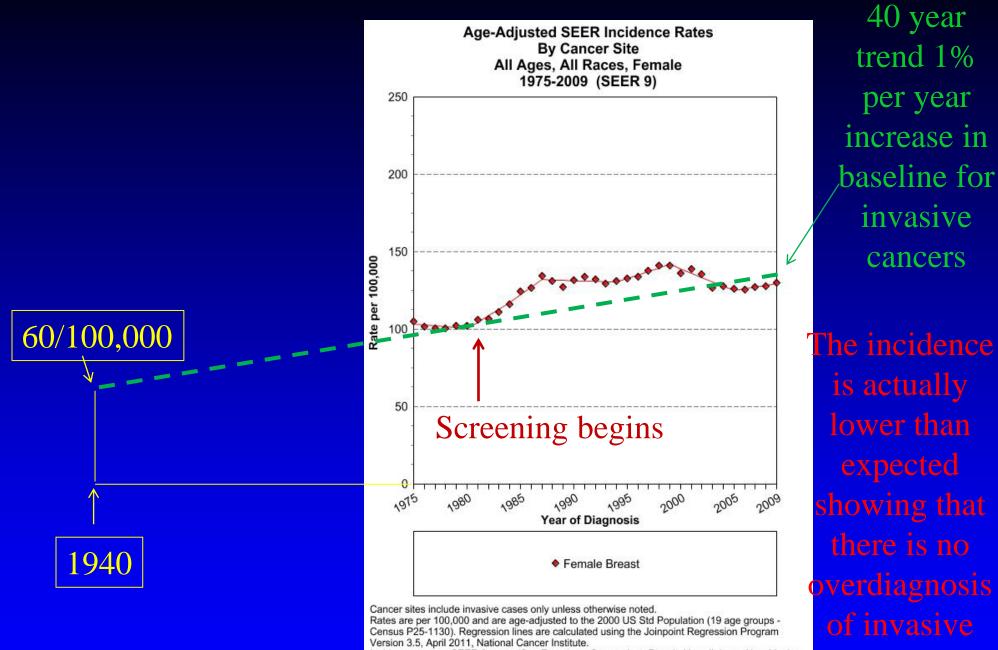
In 1940 there were 60 invasive cancers/100,000 rising to 100/100,000 in 1980. If this 1.3% per year increase continued then there should have been more than 143/100,000 in 2008 yet there were only 128/100,000.



# "OVERDIAGNOSIS" IS GROSSLY OVERESTIMATED

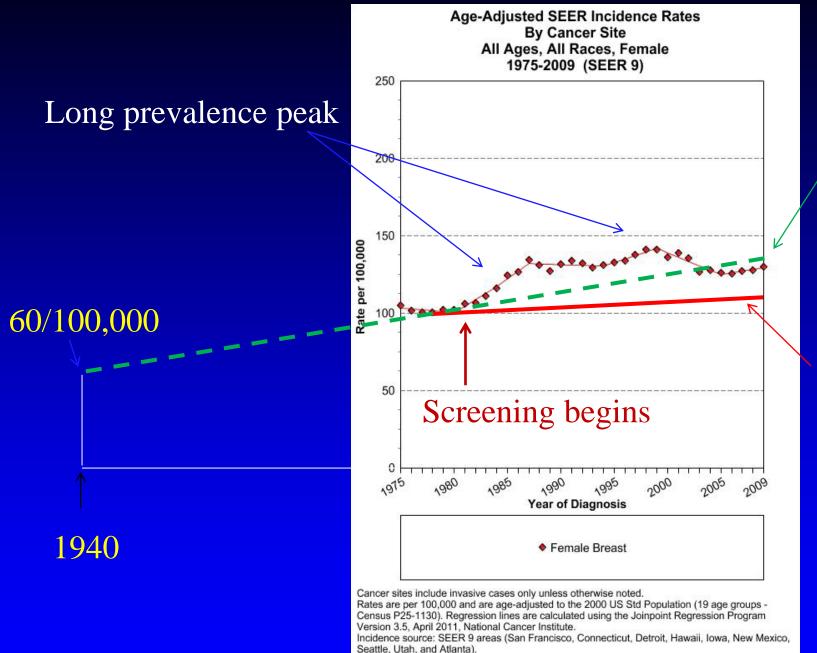
In fact, using Bleyer and Welch's approach, <u>and the correct numbers</u>, there were actually <u>fewer</u> invasive cancers in 2008 than would have been expected.





Incidence source: SEER 9 areas (San Francisco, Connecticut, Detroit, Hawaii, Iowa, New Mexico, Seattle, Utah, and Atlanta).

cancers

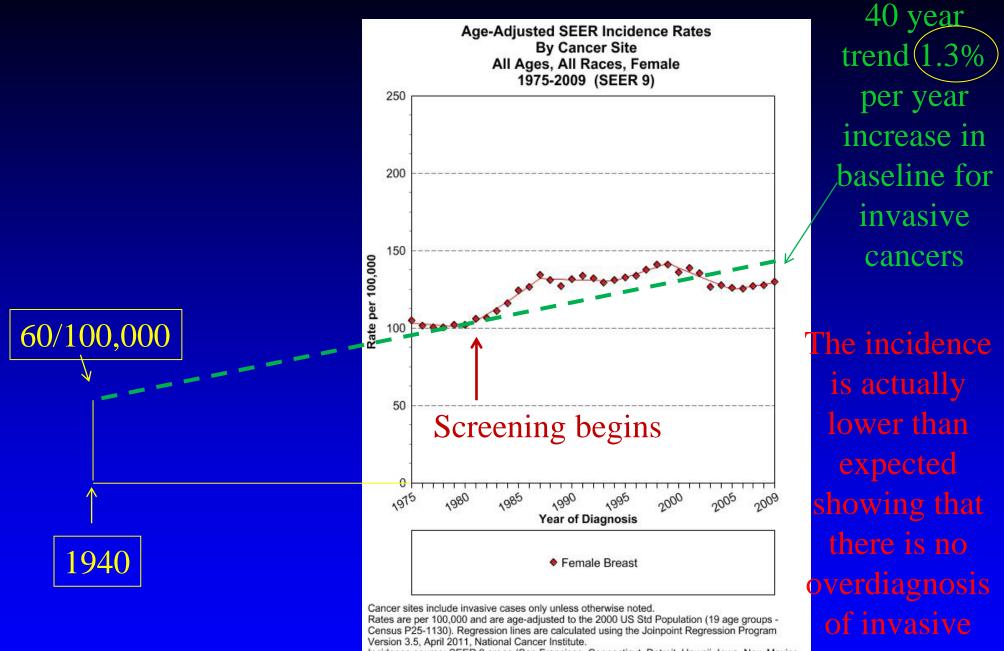


40 year trend 1% per year increase in baseline for invasive cancers Bleyer and Welch estimate 0.25% per baseline increase



# Bleyer and Welch are incorrect.





Incidence source: SEER 9 areas (San Francisco, Connecticut, Detroit, Hawaii, Iowa, New Mexico, Seattle, Utah, and Atlanta).



cancers

REMOVING DCIS FROM THE POPULATION REDUCES THE NUMBER OF FUTURE INVASIVE CANCERS

It is possible that, with longer follow-up, the removal of moderate and lower grade DCIS will further reduce the number of invasive cancer in the future.



#### **CORRECTING THE MISINFORMATION**

Bleyer and Welch claimed that there had been little if any reduction in advanced breast cancers over the time period (hence little benefit).

Actually, had they used the Connecticut Tumor Registry data

"At an APC [Annual Percentage Change] of 1.3%, late-stage breast cancer incidence decreased by 37%."

(Helvie MA, et al Reduction in late-stage breast cancer incidence in the mammography era: Implications for overdiagnosis of invasive cancer. Cancer. 2014)

More than 40 experts in breast cancer (including oncologists, surgeons, and several organizations) signed a letter to the editor of the New England Journal calling for a withdrawal of the Bleyer and Welch paper.

The NEJM refused to publish the letter.



Etzioni et al calculated that if Bleyer and Welch are correct, then American radiologists were finding invasive cancers, on average, 9 years prior to their becoming clinically evident. Even the most optimistic studies suggest a "leadtime" of 2 to at most 4 years.

#### Bleyer and Welch are wrong.

(Etzioni R, Xia J, Hubbard R, Weiss NS, Gulati R. A reality check for overdiagnosis estimates associated with breast cancer screening. J Natl Cancer Inst. 2014 Oct 31;106(12).)

There have now been 3 separate analyses that show that the conclusions of the Bleyer and Welch paper in the NEJM are incorrect.

The paper should have been withdrawn. Its conclusions should not be used to establish screening guidelines.



The "Panels" advise that delaying screening until age 50, and then biennially will reduce "overdiagnosis" (detecting "fake" cancers).



The only way that delaying screening will reduce "overdiagnosis", if it even exists, is if the "fake" cancers disappear before age 50 or between biennial screens.



No one has ever seen a mammographically detected breast cancer regress or disappear on its own.

In my informal survey of more than 2,000 breast imagers, no one has ever seen this happen.



#### BREAST CANCERS DON'T DISAPPEAR

"...among 240 untreated screen-detected invasive breast cancers, none disappeared or regressed."

## "...among 239 untreated cases of screen detected DCIS, none disappeared or regressed".

#### Total = 479

(Arleo EK, Monticciolo DL, Monsees B, McGinty G, Sickles EA.Persistent untreated screening-detected breast cancer: an argument against delaying screening or increasing the interval between screenings. J Am Coll Radiol 2017; 14:863-867.)



#### BREAST CANCERS DON'T DISAPPEAR

Since, mammographically detected, breast cancers don't disappear, delaying screening until the age of 50, or screening every two years will not reduce "overdiagnosis" [if it exists at all]. They will be there no matter when you start screening or how long you wait between screens.

(Arleo EK, Monticciolo DL, Monsees B, McGinty G, Sickles EA. Persistent untreated screening-detected breast cancer: an argument against delaying screening or increasing the interval between screenings. J Am Coll Radiol 2017; 14:863-867.)

The ONLY "harm" that delaying screening until the age of 50 or screening every two years will reduce is "recalls" for additional evaluation, most of which prove to be negative and women are told that everything is fine.



#### **REDUCE ACCESS TO SCREENING??**

Reducing access to screening will only reduce "recalls", but if women now in their thirties wait until the age of 50 and are screened every two years, as many as 100,000 will die whose lives could be saved by annual screening starting at the age of 40.

(Hendrick RE, Helvie MA. USPSTF Guidelines on Screening Mammography Recommendations: Science Ignored. Am. J. Roentgenology 2011; 196: W112 - W116.)



None of the groups seeking to reduce access have explained how many fewer recalls "balance" allowing one woman to die an avoidable death by delaying screening?!



#### MORE "ALTERNATIVE FACTS"

#### THE CLAIM THAT SCREENING DOESN'T REDUCE THE INCIDENCE OF LATE STAGE CANCERS IS

#### FALSE !



#### DECLINE IN ADVANCED DISEASE WITH SCREENING

Anderson WF, Jatoi I, Devesa SS. Assessing the impact of screening mammography: Breast cancer incidence and mortality rates in Connecticut (1943-2002). Breast Cancer Res Treat. 2006 Oct;99(3):333-40.

Tabár L, Yen AM, Wu WY, Chen SL, Chiu SY, Fann JC, Ku MM, Smith RA, Duffy SW, Chen TH. Insights from the breast cancer screening trials: how screening affects the natural history of breast cancer and implications for evaluating service screening programs. Breast J. 2015 Jan-Feb;21(1):13-20.

Yen AM, Duffy SW, Chen TH, Chen LS, Chiu SY, Fann JC, Wu WY, Su CW, Smith RA, Tabár L. Long-term incidence of breast cancer by trial arm in one county of the Swedish Two-County Trial of mammographic screening. Cancer. 2012 Dec 1;118(23):5728-32. doi: 10.1002/cncr.27580. Epub 2012 May 17

Foca F, Mancini S, Bucchi L, Puliti D, Zappa M, Naldoni C, Falcini F, Gambino ML, Piffer S, Sanoja Gonzalez ME, Stracci F, Zorzi M, Paci E; IMPACT Working Group. Decreasing incidence of latestage breast cancer after the introduction of organized mammography screening in Italy. Cancer. 2013 Jun 1;119(11):2022-8. doi: 10.1002/cncr.28014. Epub 2013 Mar 15.

Tabár L, Faberberg G, Day NE, Holmberg L. What is the optimum interval between mammographic screening examinations? An analysis based on the latest results of the Swedish two-county breast cancer screening trial. Br J Cancer. 1987 May;55(5):547-51.

Swedish Organised Service Screening Evaluation Group. Effect of mammographic service screening on stage at presentation of breast cancers in Sweden. Cancer. 2007 Jun 1;109(11):2205-12



#### DECLINE IN ADVANCED DISEASE WITH SCREENING

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#### DECLINE IN ADVANCED DISEASE WITH SCREENING

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Helvie MA, Chang JT, Hendrick RE, Banerjee M. Reduction in late-stage breast cancer incidence in the mammography era: Implications for overdiagnosis of invasive cancer. Cancer. 2014 Sep 1;120(17):2649-56. Anderson WF, Jatoi I, Devesa SS. Assessing the impact of screening mammography: breast cancer incidence and mortality rates in Connecticut (1943–2002). Breast Cancer Res Treat (2006) 99:333–340



"rates for early-stage and late stage tumors diverged in the early 1980s, also consistent with earlier detection over time.

"Breast cancer mortality rates declined 31.6%, slightly more than estimates from randomized screening trials [1, 2]. Thus, the mortality benefit of early detection and intervention seem firm."

(Anderson WF, Jatoi I, Devesa SS. Assessing the impact of screening mammography: breast cancer incidence and mortality rates in Connecticut (1943–2002). Breast Cancer Res Treat (2006) 99:333–340)

#### EPIDEMIOLOGY

Assessing the impact of screening mammography: breast cancer incidence and mortality rates in Connecticut (1943–2002)

William F. Anderson · Ismail Jatoi · Susan S. Devesa

REGIONAL AND DISTANT DISEASE DECLINED AS MORE AND MORE WOMEN PARTICIPATED IN SCREENING

reast Cancer Res Treat (2006) 99:333-340

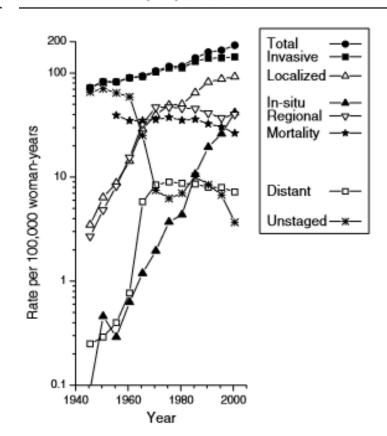
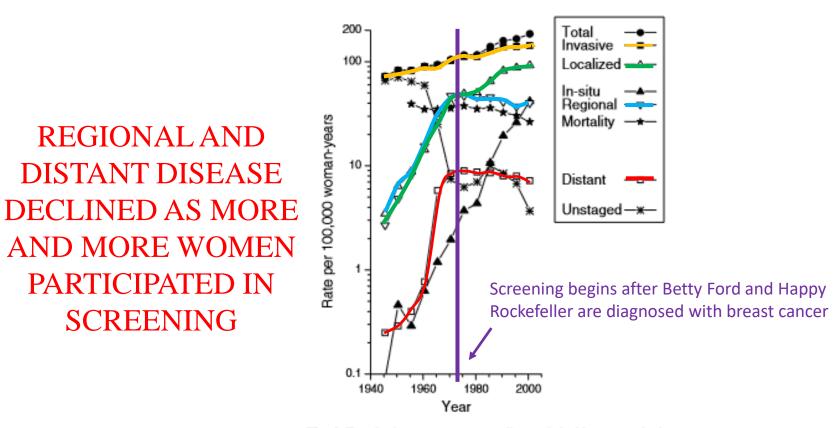


Fig. 1 Female breast cancer mortality and incidence trends in Connecticut by stage and twelve 5-year time periods of diagnosis (1943–1947 to 1998–2002). Note: "Total' includes *in-situ* and invasive breast cancers



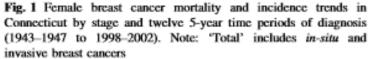


Breast Cancer Res Treat (2006) 99:333-340

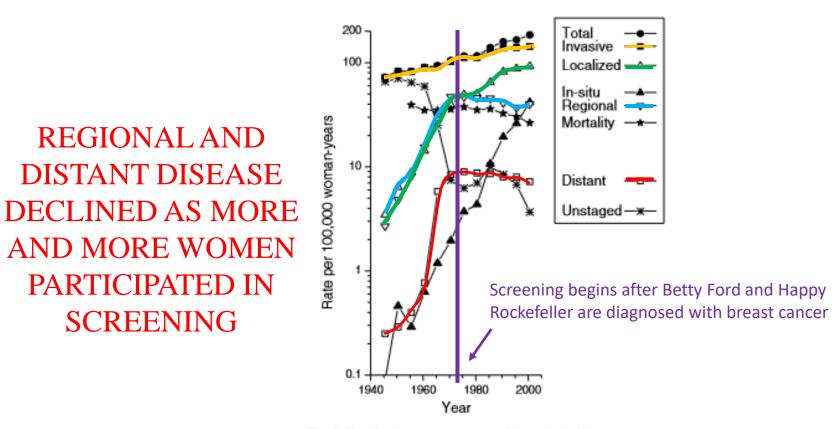
**REGIONAL AND** 

PARTICIPATED IN

**SCREENING** 





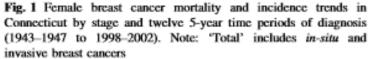


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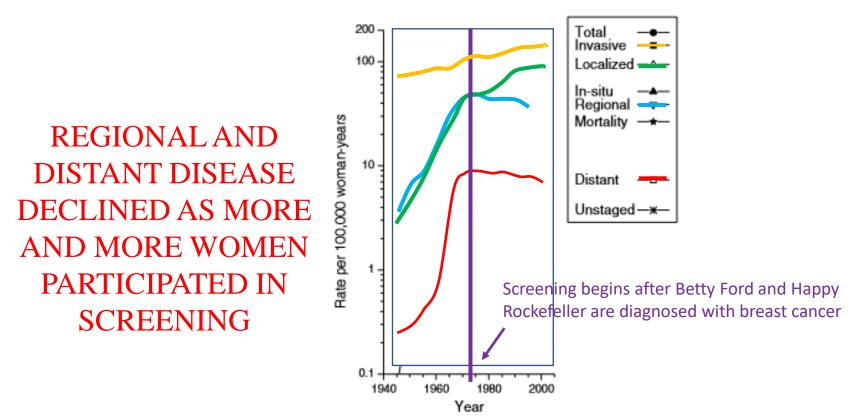
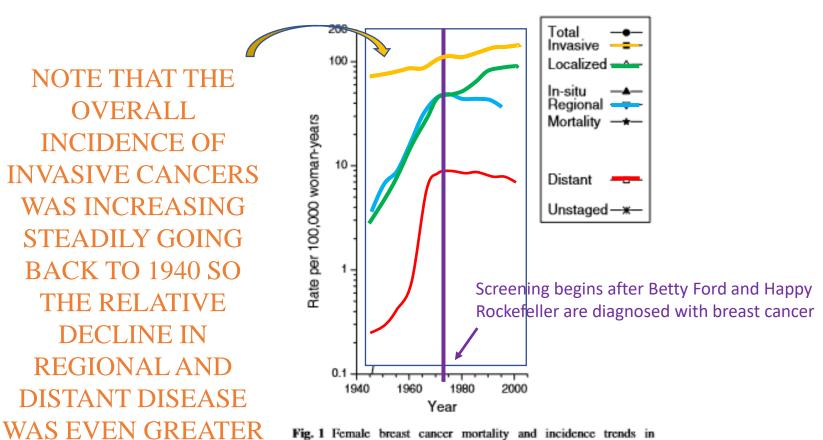


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Breast Cancer Res Treat (2006) 99:333-340

**OVERALL** 

**INCIDENCE OF** 

THE RELATIVE

**DECLINE IN** 

Fig. 1 Female breast cancer mortality and incidence trends in Connecticut by stage and twelve 5-year time periods of diagnosis (1943-1947 to 1998-2002). Note: 'Total' includes in-situ and invasive breast cancers



#### MORE "NONSCIENCE" FROM THE NEW ENGLAND JOURNAL OF MEDICINE

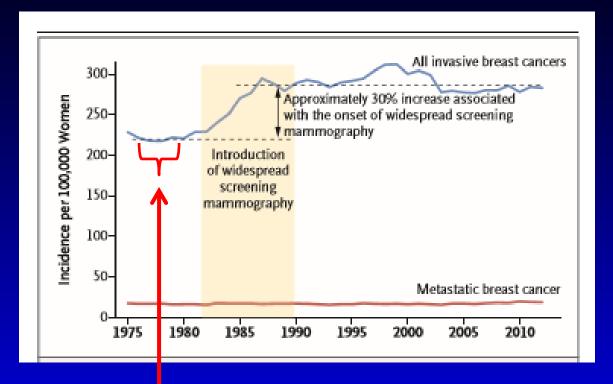
The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

Breast-Cancer Tumor Size, Overdiagnosis, and Mammography Screening Effectiveness

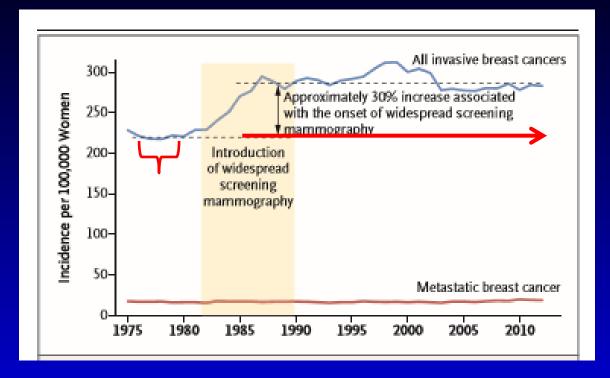
H. Gilbert Welch, M.D., M.P.H., Philip C. Prorok, Ph.D., A. James O'Malley, Ph.D., and Barnett S. Kramer, M.D., M.P.H.

The authors claimed that the shift to smaller cancers related to mammography screening was, predominantly "overdiagnosed" cancers since the rate of metastatic disease was only slightly reduced.



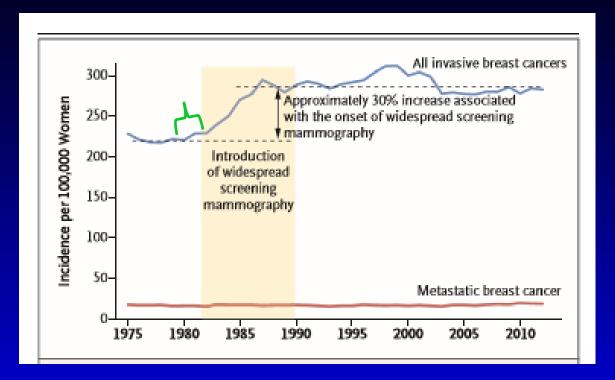
The authors, incorrectly, decided that the underlying incidence in the absence of screening was defined by the period 1975-1979.





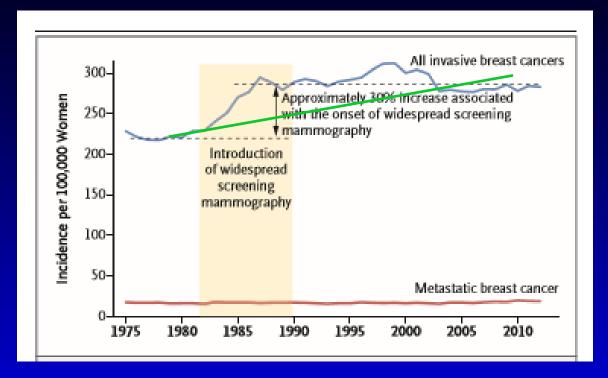
They claimed that in the absence of screening the incidence of cancer would have been a flat line and would have been the same in 2012 as in 1977 (red line).



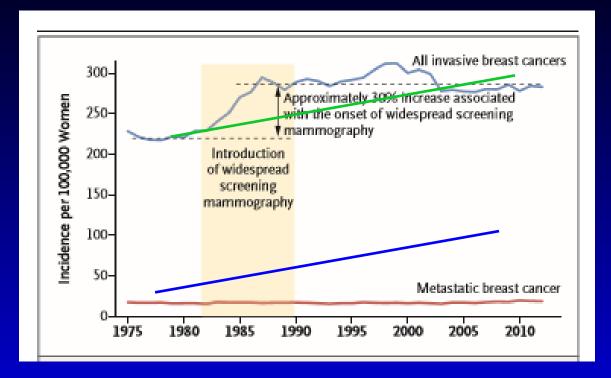


The actual increase in underlying incidence was 1 - 1.3% per year which is similar to what is seen in the prescreening period from 1979-1982 (green).





If the correct extrapolation is used there is no overdiagnosis (green line). There are actually fewer cancers than expected, likely due to the removal of DCIS lesions preventing them from becoming invasive.



If the correct extrapolation is used the rate of metastatic disease should have increased at the same rate as the other invasive cancers (blue line). This means that the rate of metastatic disease has declined dramatically (red line).



#### MORE "NONSCIENCE" FROM THE NEW ENGLAND JOURNAL OF MEDICINE

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Not only was the conclusion of this paper scientifically unsupportable, but the lead author left Dartmouth having been found guilty of plagiarism!



#### MORE "NONSCIENCE" FROM THE NEW ENGLAND JOURNAL OF MEDICINE



In 2012 Welch argued that the baseline incidence of breast cancer would have increased by 0.25%-0.5% per year. In this paper for the same time period he claimed it was 0.0%!

Same data, same journal different numbers!

Great peer review at the NEJM!





If you don't know what you are talking about it might be a good idea to not talk about it!



One of the more recent pseudo issues raised by those seeking to reduce access to screening is the claim that the 30% reduction in breast cancer deaths in the RCT's did not, significantly, reduce deaths from all causes.



In treatment trials it is important to look at deaths from all causes because your treatment might reduce breast cancer deaths, but cause deaths from other problems

eg. Breast radiation therapy caused deaths from heart damage.



### "ALL CAUSE" MORTALITY THE LATEST "NONSCIENCE"

However, in treatment trials, since everyone has breast cancer, the vast majority of deaths will be due to breast cancer and not from other "all causes" so that a reduction in breast cancer deaths is likely to reduce total ("all cause") deaths.



### "ALL CAUSE" MORTALITY THE LATEST "NONSCIENCE"

In screening trials that evaluate a normal population, a very small number of women develop breast cancer, and an even smaller number die. Most deaths in the trial will be due to causes other than breast cancer.



#### "ALL CAUSE" MORTALITY IS THE LATEST "NONSCIENCE"

Each year only, approximately, 3% of deaths in the general population are due to breast cancer. A 30% reduction in breast cancer deaths will only reduce, "all cause", total mortality by 1%. You would need approximately 2.5 million women in a trial to show this reduction as significant.

(Tabar L, Duffy SW, Yen MF, Warwick J, Vitak B, Chen HH, Smith RA. All-cause mortality among breast cancer patients in a screening trial: support for breast cancer mortality as an end point. J Med Screen. 2002;9(4):159-62.)

#### "ALL CAUSE" MORTALITY THE LATEST "NONSCIENCE"

If you look at women diagnosed with breast cancer in RCT's (so that they are like treatment trials), reducing breast cancer deaths, significantly, reduces all cause mortality.

(Tabar L, Duffy SW, Yen MF, Warwick J, Vitak B, Chen HH, Smith RA. All-cause mortality among breast cancer patients in a screening trial: support for breast cancer mortality as an end point. J Med Screen. 2002;9(4):159-62.)

#### BREAST CANCER SCREENING

# !! CAUTION !! "RISK BASED" SCREENING And "VALUE BASED" SCREENING

are stealth efforts to reduce access to screening.



There are two interpretations of

#### "RISK BASED SCREENING".

 The ACR and the SBI advise that all women be screened annually starting at the age of 40 with very high risk women beginning at younger ages depending on risk. Women with a lifetime risk of 25% or more should consider alternating mammography with MRI every 6 months.



#### CAUTION!!

#### "RISK BASED" SCREENING

2. Groups want to limit screening to women who will develop breast ! (as if no one ever thought of doing that before). They want to ONLY screen high risk women.



#### CAUTION!!

#### "RISK BASED" SCREENING

 Only 10% of women who are diagnosed with breast cancer each year have a BRCA1 or 2 mutation.
 Another 15% have a family history or other elevator of risk.



## !!CAUTION!! "RISK BASED" SCREENING 2:

It is "pie in the sky" to suggest that screening can be tailored based on risk.

1. The randomized, controlled trials were not stratified by risk so there is no proof that screening only high risk women will save any lives.



## !!CAUTION!! "RISK BASED" SCREENING 2:

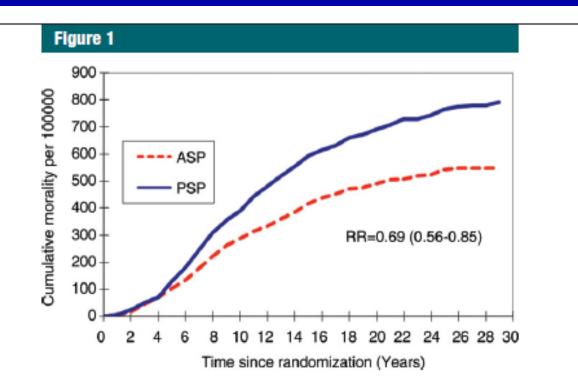
It is "pie in the sky" to suggest that screening can be tailored based on risk.

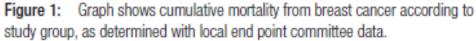
If we only screen high risk women,
 of women who develop breast
 cancer each year will not benefit from
 early detection.



#### FACT:

Screening has, consistently, shown a decrease in breast cancer deaths for all women of approximately 30%.



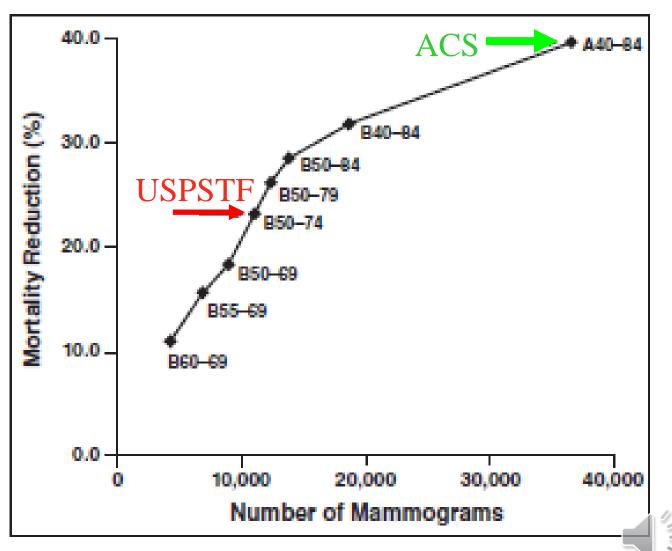


Tabár L, Vitak B, Chen TH, Yen AM, Cohen A, Tot T, Chiu SY, Chen SL, Fann JC, Rosell J, Fohlin H, Smith RA, Duffy SW. Swedish two-county trial: impact of mammographic screening on breast cancer mortality during 3 decades. Radiology. 2011 Sep;260(3):658-63.



#### USPSTF SHOWS THAT MOST LIVES ARE SAVED BY ANNUAL SCREENING BEGINNING AT 40

Fig. 1—Percentage mortality reduction from various screening strategies. Note that annual (A) screening from ages 40–84 years (A40–84, *solid arrow*) is estimated to have 71% greater mortality benefit than biennial (B) screening from ages 50–74 years (B50–74, *dashed arrow*). Number of mammograms shown on horizontal axis is per 1,000 women screened. Data shown are mean values of six models from [6].



#### **BREAST CANCER SCREENING**

#### The Bottom Line

Most women who develop breast cancer are not at increased risk.

All women are at risk and annual screening, beginning at the age of 40, should be encouraged for all women.



#### ACR 2017 SCREENING GUIDELINES

"Women should be helped to understand the risks of screening;

weighing benefits and risks *should* be done by women, not for women."

(Monticciolo DL, Newell MS, Hendrick RE, Helvie MA, Moy L, Monsees B, Kopans DB, Eby PR, Sickles EA. Breast Cancer Screening for Average-Risk Women: Recommendations From the ACR Commission on Breast Imaging. J Am Coll Radiol. 2017 Sep;14(9):1137-1143.)