

Breast Cancer Screening: Consensus and Controversies

Michael Morris, MD

Do you have a family member or friend with a history of breast cancer?

1. Yes
2. No

Do you have a family member or friend with a history of breast cancer?

1. Yes
2. No

Common Types of Cancer	Estimated New Cases 2014	Estimated Deaths 2014
1. Prostate Cancer	233,000	29,480
2. Breast Cancer (Female)	232,670	40,090
3. Lung and Bronchus Cancer	224,210	152,260
4. Colon and Rectum Cancer	136,830	50,310
5. Melanoma of the Skin	76,100	9,710
6. Bladder Cancer	74,690	15,580
7. Non-Hodgkin Lymphoma	70,800	18,990
8. Kidney and Renal Pelvis Cancer	63,920	13,860
9. Thyroid Cancer	63,080	1,890
10. Endometrial Cancer	52,630	8,590

Breast cancer represents 14.0% of all new cancer cases in the U.S.

14.0%

<http://seer.cancer.gov/statfacts/html/breast.html>

What is the probability of a 50 year old women having breast cancer in the next 10 years?

1. 2%
2. 12%
3. 20%

What is the probability of a 50 year old women having breast cancer in the next 10 years?

1. 2%
2. 12%
3. 20%

<http://seer.cancer.gov/canstat/animat>

What is the probability of a 50 year old women having breast cancer in the next 10 years?

1. 2%
2. 12%
3. 20%

	Median (interquartile range)	
	All subjects	Insurers
Probability of developing breast cancer within the next 10 years		
Personal, %	20.8 (16.0-30.0)	
Calculated, %	2.7 (2.0-2.8)	
Ratio	5.9 (3.1-13.8)	4.8† 9.7

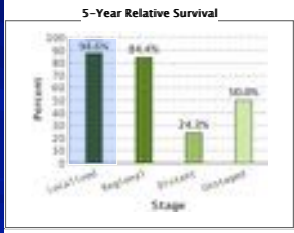
J Natl Cancer Inst. 1995;87(10):720-31

What is the risk of dying from early stage breast cancer?

1. 1%
2. 10%
3. 20%
4. 30%

What is the risk of dying from early stage breast cancer?

1. 1%
2. 10%
3. 20%
4. 30%



<http://seer.cancer.gov/statfacts/html/breast.html>

What is the risk of dying from early stage breast cancer?

1. 1%
2. 10%
3. 20%
4. 30%

	DCIS (N = 49)	IBC (N = 392)
How likely is it that you will die of breast cancer?		
Mean score	1.8	3.9
1 (Not likely at all)	45/49 (91%)	219/392 (56%)
2 (Somewhat unlikely)	3/49 (6%)	161/392 (41%)
3 (Somewhat likely)	0/49 (0%)	20/392 (5%)
4 (Very likely)	0/49 (0%)	52/392 (13%)
How likely is it that you will die of something other than breast cancer?		
Mean score	1.6	3.5
1 (Not likely at all)	11/49 (22%)	26/392 (7%)
2 (Somewhat unlikely)	32/49 (65%)	241/392 (61%)
3 (Somewhat likely)	0/49 (0%)	76/392 (19%)
4 (Very likely)	0/49 (0%)	29/392 (7%)

*DCIS: ductal carcinoma in situ; IBC: early stage invasive breast cancer

Breast Cancer Res Treat. 2003;77(3):285-93

Breast Cancer Screening

- Where there is consensus I say so
- Where there is controversy I say so
- Abolishing Mammography Screening Programs? A View from the Swiss Medical Board
- Make your own informed decision



My Advice:

- Stick with recommendations of major medical societies
 - ACS (American Cancer Society)
 - NCI (National Cancer Institute)
 - USPSTF (US Preventative Services Task Force)
 - ACP (American College of Physicians)
 - AAFP (American Academy of Family Physicians)
 - ACR (American College of Radiology)
 - ACOG (American College of Obstetrics & Gynecology)

Methods of Breast Cancer Screening

- Physical Exam
 - Breast self exam
 - Clinical breast exam
- Imaging
 - Mammography
 - MRI
 - Ultrasound
 - Tomosynthesis, molecular breast imaging, contrast mammography
- Blood biomarkers

Definitions

- Screening Exam:
 - Look for breast cancer in asymptomatic women
- Diagnostic Exam:
 - Referred for specific concern
 - Palpable lump, focal pain, nipple discharge (*clear, bloody*)

Definitions

- Screening Exam:
 - Look for breast cancer in asymptomatic women
- Diagnostic Exam:
 - Referred for specific concern
 - Palpable lump, focal pain, nipple discharge (*clear, bloody*)

Case #1

- 41 y/o woman comes for her annual exam
 - No PMH; No FH cancer
 - Occasional EtOH; non-smoker
- You perform routine exam, including clinical breast exam

Physical Exam Screening

<p><u>CLINICAL BREAST EXAM</u></p> <ul style="list-style-type: none"> ■ Yes: <ul style="list-style-type: none"> • ACS, NCI, ACOG ■ Insufficient data: <ul style="list-style-type: none"> • USPSTF, AAFP 	<p><u>BREAST SELF EXAM</u></p> <ul style="list-style-type: none"> ■ Yes: <ul style="list-style-type: none"> • ACOG ■ No: <ul style="list-style-type: none"> • USPSTF, AAFP, ACS, NCI
--	---


Would you refer this patient for a screening mammogram?

- 41 y/o woman comes for her annual exam
 - No PMH; No FH cancer
 - Occasional EtOH; non-smoker
- You perform routine exam, including clinical breast exam

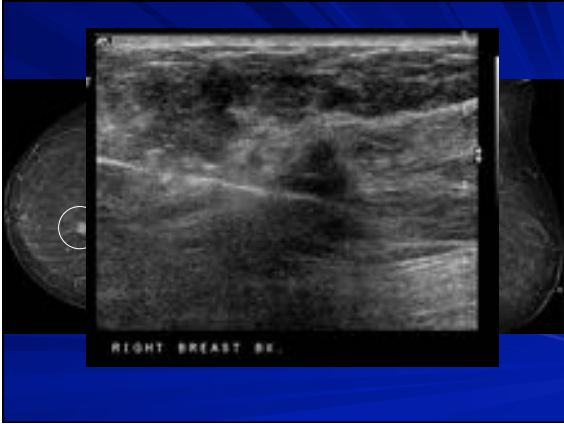
1. Yes
2. No

Consensus

- Only screening method to reduce mortality in RCT
 - Ages 40-74
- Overall ~20% mortality reduction
 - Sensitivity 84.9%
 - Specificity 90.3%
- Yes: ACS, NCI, USPSTF, AAFP, ACOG, ACR, ACP
- No: 0



Lancet. 2012; 380:1778-86
<http://breastscreening.cancer.gov/data>



Benefits of Screening vs "Harms" of Screening

Benefits of Screening

- 20% mortality reduction*
 - If 1000 women screened
 - Absolute decrease 5 → 4

→

- Very big benefits, few people

"Harms" of Screening

Ann Intern Med. 2009;151:727-37
J Med Screen 2012;19 s1:42-56

Benefits of Screening

- 20% mortality reduction*
 - If 1000 women screened
 - Absolute decrease 5 → 4

→

- Very big benefits, few people

"Harms" of Screening

- False Positives
 - ~10% recalled: ~1% biopsy
 - Anxiety, morbidity, cost
- High NNT when younger
 - 40-50: 1/1900
 - 60-70: 1/377

→

- Small harms, many people

Ann Intern Med. 2009;151:727-37
J Med Screen 2012;19 s1:42-56

Benefits of Screening

- 20% mortality reduction*
 - Very big benefits, few people

"Harms" of Screening

- False Positives
 - Small harms, many people
- High NNT when younger
- Overdiagnosis
 - ~1-20% women with cancer Rx unnecessarily
 - Big harms, very few people

Ann Intern Med. 2009;151:727-37
J Med Screen 2012;19 s1:42-56

When to start screening?
• Age 40 or 50

Screening frequency?
• Yearly or biennial

When to stop?
• Age 75, 80, 85

Very big benefits, few people

Small harms, many people
Big harms, very few people

▲

- When to start screening?
 - Age 40 or 50
- Screening frequency?
 - Yearly or biennial
- When to stop?
 - Age 75, 80, 85

Very big benefits,
few people

Small harms,
many people
Big harms,
very few people

JAMA 2014;311(13):1327-35

Organization and Year of Evidence	Recommendation Regarding Mammography Screening
International Breast Cancer Screening Program, 1992 ¹	Screening mammography every 2 y for women between ages 50 y and 69 y
US Preventive Services Task Force, 2009 ²	Biennial screening mammography for women between ages 50 y and 74 y The decision to start regular, biennial screening mammography before age 40 y should be an individual one and take into account patient's values, including the patient's wishes regarding specific tests and tests.
National Health Service Breast Screening Program (United Kingdom), 2007 ³	Screening mammography every 3 y for women aged 47-73 y
Canadian Task Force on Preventive Health Care, 2011 ⁴	Active screening mammography for women aged 50-74 y
National Cancer Institute (United States), 2012 ⁵	Screening mammography every 1 to 2 y in women 40-74 y
American Cancer Society (United States), 2012 ⁶	Yearly screening and starting at age 40 y

Very big benefits,
few people

Small harms,
many people
Big harms,
very few people

JAMA 2014;311(13):1327-35

Organization and Year of Evidence	Recommendation Regarding Mammography Screening
International Breast Cancer Screening Program, 1992 ¹	Screening mammography every 2 y for women between ages 50 y and 69 y
US Preventive Services Task Force, 2009 ²	Biennial screening mammography for women between ages 50 y and 74 y The decision to start regular, biennial screening mammography before age 40 y should be an individual one and take into account patient's values, including the patient's wishes regarding specific tests and tests.
National Health Service Breast Screening Program (United Kingdom), 2007 ³	Screening mammography every 3 y for women aged 47-73 y
Canadian Task Force on Preventive Health Care, 2011 ⁴	Active screening mammography for women aged 50-74 y
National Cancer Institute (United States), 2012 ⁵	Screening mammography every 1 to 2 y in women 40-74 y
American Cancer Society (United States), 2012 ⁶	Yearly screening and starting at age 40 y

Very big benefits,
few people

Small harms,
many people
Big harms,
very few people

JAMA 2014;311(13):1327-35

Organization and Year of Evidence	Recommendation Regarding Mammography Screening
American Cancer Society (United States), 2012 ⁶	Yearly screening and starting at age 40 y
National Health Service Breast Screening Program (United Kingdom), 2007 ³	Screening mammography every 3 y for women aged 47-73 y

US Breast Cancer Mortality: 22/100,000

UK Breast Cancer Mortality: 24/100,000

Very big benefits,
few people

Small harms,
many people
Big harms,
very few people

JAMA 2014;311(13):1327-35

When to Start Screening? (average risk women)

Age 40

- RCT trials start age 40
- 15% mortality reduction* in women 40-49
 - *relative decrease
- Yes: ACS, NCI, ACR, ACOG

Wait until age 50

- Breast Ca incidence is lower (high NNT)
- Harms outweigh benefits
 - Increased stress from false positive mammogram
 - Increased biopsies
- Yes: USPSTF, AAFP

Ann Intern Med. 2007;146(7):511-5

When to Stop Screening?

- RCT data stop @ age 74
- Mortality benefit mammography 7-10 yrs
- Consider stop screening if:
 - Life expectancy <7-10yrs
 - Not willing to undergo f/u if abnrml mammo
- ACS, ACR, ACOG, AAFP

JACR 2010;7:18-27

Case #2

- 35 y/o women, new patient visit
 - Healthy
 - Mom breast ca age 45
 - No other FH ca
 - G2P2, first birth age 32

Should you refer this patient for a screening mammogram?

- 35 y/o women, new patient visit
 - Healthy
 - Mom breast ca age 45
 - No other FH ca
 - G2P2, first birth age 32

1. Yes
2. No
3. Not Sure

Should you refer this patient for a breast MRI?

- 35 y/o women, new patient visit
 - Healthy
 - Mom breast ca age 45
 - No other FH ca
 - G2P2, first birth age 32

1. Yes
2. No
3. Not Sure


Case #2

- 35 y/o women, new patient visit
 - Healthy
 - Mom breast ca age 45
 - No other FH ca
 - G2P2, first birth age 32

■ Is she at high risk for breast cancer?

Who is High Risk?

- ACS expert panel review of evidence
- High risk = 20-25% lifetime risk breast ca
 - Supplement mammography screening with MRI



CA Cancer J Clin 2007;57:75-89

When to Start High Risk Screening?


- BRCA or chest wall radiation
 - 10 years after chest wall radiation
 - Not before age 25
- Family history
 - 10 years before onset of cancer in relative
- MRI does not replace mammography

JACR 2010;7:18-27
J Clin Oncol. 2010; 28(9):1450-7

Who is High Risk?

20-25% Lifetime Risk:

- BRCA
 - Untested 1st degree relative
- Chest wall radiation 10-30 year old
- Li Fraumeni, Cowden
- Calculated lifetime risk by models (PMH, FH)
 - Gail
 - Tyrer-Cuzick
 - BRCAPRO



www.cancer.gov/bcrisktool/
CA Cancer J Clin 2007;57:75-89

Who is (Not) High Risk?

20-25% Lifetime Risk:

- BRCA
 - Untested 1st degree relative
- Chest wall radiation 10-30 year old
- Li Fraumeni, Cowden
- Calculated lifetime risk by models (PMH, FH)
 - Gail
 - Tyrer-Cuzick
 - BRCAPRO

Not recommended (<20% Lifetime Risk)

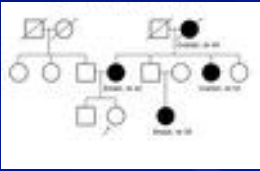
- Personal history breast cancer
- Increased breast density on mammography

www.cancer.gov/bcrisktool/
CA Cancer J Clin 2007;57:75-89

Who should be referred to genetic counseling?

NCCN Guidelines:

- 2 or more breast ca on same side of family
- Breast ca in 1st or 2nd degree relative <45 y/o
- Male breast ca
- Ovarian ca
- Gene mutation in susceptibility gene

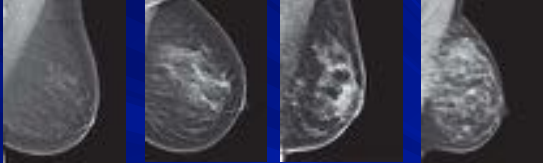


J Natl Compr Canc Netw. 2009;7(10):1060-96

Case #3

- 53 y/o woman calls your office
 - Mammogram report states: "dense breasts, which can lower the sensitivity of mammography"
- "What does this mean?"
- "Do I need additional testing?"

Breast Density

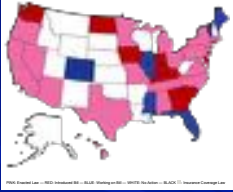



Entirely fat (~10%)	Scattered fibroglandular (~40%)	Heterogeneously dense (~40%)	Extremely dense (~10%)
Sens: 88.2 Spec: 96.5	Sens: 82.1 Spec: 93	Sens: 68.9 Spec: 90.8	Sens: 62.2 Spec: 89.9

Ann Intern Med 2003;138:168-175

Breast Density – Emerging Controversy

- Higher breast density
 - lower sensitivity
 - Independent risk factor? (~2-4x)
- Mandated reporting

■ Your mammogram indicates that you have dense breast tissue. Dense breast tissue is common and is found in 50% of women. However, dense breast tissue can make it difficult to detect cancers in the breast by mammography and may also be associated with an increased risk of breast cancer. This information is being provided to raise your awareness and to encourage you to discuss with your health care providers your dense breast tissue and other breast cancer risk factors. Together, you and your physician can decide if additional screening options are right for you.

Summary

<p><u>Consensus</u></p> <ul style="list-style-type: none"> ■ Clinical breast exam ■ Screening mammography in average risk women ■ Screening mammography & MRI in high risk women 	<p><u>Controversy</u></p> <ul style="list-style-type: none"> ■ When to start? ■ How often? ■ When to stop? ■ Breast density?
--	---

Very big benefits,
few people

Small harms,
many people
Big harms,
very few people

Future

- Standardization → Individualization
 - Screening based on density
 - Fatty breasts screened less often?
 - New technologies
 - Tomosynthesis (3D mammography)
 - Automated breast ultrasound
 - Molecular imaging & genomics