Canadian Task Force on Preventive Health Care

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Appendix 1: PRISMA Checklist

Section/topic	#	Checklist item	Reported on page #	
TITLE	TITLE			
Title	1	Identify the report as a systematic review, meta-analysis, or both.	10 (Modified overview and update)	
ABSTRACT				
Structured summary	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.		
INTRODUCTION				
Rationale	3	Describe the rationale for the review in the context of what is already known.		
Objectives	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).		
METHODS	1			
Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number.		
Eligibility criteria	6	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.		
Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.		
Search	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.		
Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).		
Data collection process	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.		

Data items	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	
Risk of bias in individual studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	
Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	
Synthesis of results	14	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., I²) for each meta-analysis.	

Page 1 of 2

Section/topic	#	Checklist item	Reported on page #
Risk of bias across studies	15	Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).	
Additional analyses	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	
RESULTS	•		
Study selection	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	
Study characteristics	18	or each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and rovide the citations.	
Risk of bias within studies	19	resent data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	
Results of individual studies	dies 20 For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.		
Synthesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency.	
Risk of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).	
Additional analysis	23	23 Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).	
DISCUSSION			
Summary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	

Limitations	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete identified research, reporting bias).		
Conclusions 26 Provide a general interpretation of the results in the context of other evidence, and implications for future research.			
FUNDING			
Funding	27	Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the systematic review.	

From: Moher D, Liberati A, Tetzlaff J, Altman DG, The PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. PLoS Med 6(7): e1000097. doi:10.1371/journal.pmed1000097

For more information, visit: www.prisma-statement.org.

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Appendix 2- Search Strategy (Updated Search)

Final Strategies 2017 Jan 4

EFFECTIVENESS

MEDLINE

Database: Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE(R) Daily and Ovid MEDLINE(R) <1946 to Present> Search Strategy:

.....

- 1 exp Breast Neoplasms/ (287642)
- 2 ((breast* or mamma or mammar*) adj3 (cancer* or carcinoid* or carcinoma* or carcinogen* or adenocarcinoma* or adeno-carcinoma* or malignan* or neoplasia* or neoplasm* or sarcoma* or tumour* or tumor*)).tw,kw. (331761)
- 3 exp Carcinoma, Intraductal, Noninfiltrating/ (10262)
- 4 intraductal carcinoma*.tw,kw. (922)
- 5 (ductal carcinoma in situ or DCIS).tw,kw. (7495)
- 6 or/1-5 [BREAST CANCER] (395424)
- 7 exp Breast Neoplasms/di, pc (47714)
- 8 exp Mass Screening/ (124871)
- 9 screen*.tw,kw. (660022)
- 10 "Early Detection of Cancer"/ (18397)
- 11 ((early or earlier or earliest) adj3 (detect* or diagnos* or identif* or recogni*)).tw,kw.(225345)
- 12 exp Self-Examination/ (2485)
- 13 ((self-exam* or self-detect* or self-screen*) adj5 (breast\$1 or mamma or mammary or nipple\$1)).tw,kw. (2050)
- 14 Physical Examination/ (40906)
- 15 (exam* adj5 (breast? or mamma or mammar* or nipple?)).tw,kw. (15055)
- 16 exp Breast Neoplasms/ra (16756)
- 17 exp Mammography/ (32349)
- 18 (mammograph* or mammogram*).tw,kw. (33182)
- 19 exp Magnetic Resonance Imaging/ (415717)
- 20 (fMRI or fMRIs or MRI or MRIs or NMRI or NMRIs or MR imaging or NMR imaging or magnetic resonance imag* or magnetic resonance tomograph* or MR tomograph*).tw,kw. (380636)
- 21 (chemical shift imaging or proton spin tomograph* or zeugmatograph*).tw,kw. (1076)
- 22 exp Breast Neoplasms/us (4023)
- 23 (ultrasound* or ultrason* or echograph* or echomammogra* or echo-mammogra* or echotomograph* or echo-tomograph* or sonograph*).tw,kw. (382288)
- 24 Imaging, Three-Dimensional/ (64456)
- 25 ((3D or "3-D") adj3 imag*).tw,kw. (17743)
- 26 (("3" or three) adj dimension* adj3 imag*).tw,kw. (15527)
- 27 tomosynthes*.tw,kw. (1236)
- 28 or/7-27 (1875233)

```
6 and 28 [BREAST CANCER SCREENING] (102429)
    Male/ not (Female/ and Male/) (2788208)
31
    29 not 30 [MALE-ONLY REMOVED] (100934)
32
    exp Infant/ not (exp Adult/ and exp Infant/) (838449)
    exp Child/ not (exp Adult/ and exp Child/) (1197384)
33
34
    Adolescent/ not (exp Adult/ and Adolescent/) (595774)
35
    or/32-34 (1865046)
    31 not 35 [CHILD-ONLY REMOVED] (100454)
36
    exp Animals/ not (exp Animals/ and Humans/) (4850259)
37
    36 not 37 [ANIMAL-ONLY REMOVED] (98888)
38
39
    (comment or editorial or news or newspaper article).pt. (1254980)
40
    (letter not (letter and randomized controlled trial)).pt. (1008588)
    38 not (39 or 40) [OPINION PIECES REMOVED] (91963)
41
    (201410* or 201411* or 201412* or 2015* or 2016* or 2017*).dc. (3219115)
42
43
    41 and 42 [UPDATE PERIOD] (13074)
    (controlled clinical trial or randomized controlled trial or pragmatic clinical trial).pt. (600336)
44
45
    clinical trials as topic.sh. (197690)
46
    (randomi#ed or randomly or RCT$1 or placebo*).tw. (882744)
47
    ((singl* or doubl* or trebl* or tripl*) adj (mask* or blind* or dumm*)).tw. (167447)
48 trial.ti. (201433)
    or/44-48 (1273229)
49
50 43 and 49 [RCTS] (1017)
    remove duplicates from 50 [RCTS - DUPLICATES REMOVED] (738)
51
********
Cochrane Library
Search Name: CTFPHC - Breast Cancer Screening - All Modalities
Date Run:
              04/01/17 17:35:49.798
Description:
              2017 Jan 4 (OHRI) - Oct 2014-present - FINAL
ID
       Search Hits
#1
       [mh "Breast Neoplasms"]
                                     9949
#2
       ((breast* or mamma or mammar*) near/3 (cancer* or carcinoid* or carcinoma* or carcinogen*
or adenocarcinoma* or adeno-carcinoma* or malignan* or neoplasia* or neoplasm* or sarcoma* or
tumour* or tumor*)):ti,ab,kw 22627
#3
       [mh "Carcinoma, Intraductal, Noninfiltrating"] 118
#4
       (intraductal next carcinoma*):ti,ab,kw 181
#5
       ("ductal carcinoma in situ" or DCIS):ti,ab,kw
                                                    302
#6
       {or #1-#5}
                      22683
       [mh "Breast Neoplasms"/DI,PC] 1459
#7
#8
       [mh "Mass Screening"] 5540
#9
       screen*:ti,ab,kw
                              29461
       [mh "Early Detection of Cancer"]
#10
                                             898
       ((early or earlier or earliest) near/3 (detect* or diagnos* or identif* or recogni*)):ti,ab,kw
#11
       5661
```

#12

[mh Self-Examination] 202

```
#13
       ((self next (exam* or detect* or screen*)) near/5 (breast* or mamma or mammary or
nipple*)):ti,ab,kw
                      208
#14
       [mh ^"Physical Examination"]
                                    913
#15
       (exam* near/5 (breast* or mamma or mammar* or nipple*)) .tw,kw.
                                                                          2
#16
       [mh "Breast Neoplasms"/ra]
                                     380
#17
       [mh Mammography]
                             1033
#18
       (mammograph* or mammogram*):ti,ab,kw
                                                    1859
#19
       [mh "Magnetic Resonance Imaging"]
                                            7076
#20
       (fMRI or fMRIs or MRI or MRIs or NMRI or NMRIs or "MR imaging" or "NMR imaging" or
("magnetic resonance" next imaging) or ("magnetic resonance" next tomograph*) or (MR next
tomograph*)):ti,ab,kw 14833
       ("chemical shift imaging" or ("proton spin" next tomograph*) or zeugmatograph*):ti,ab,kw
#21
       20
#22
       [mh "Breast Neoplasms"/US]
                                     86
#23
       (ultrasound* or ultrason* or echograph* or echomammogra* or echo-mammogra* or
echotomograph* or echo-tomograph* or sonograph*):ti,ab,kw 21358
#24
       [mh "Imaging, Three-Dimensional"]
                                            1022
#25
       ((3D or "3-D") near/3 imag*):ti,ab,kw
                                            338
#26
       (((3 or three) next dimension*) near/3 imag*):ti,ab,kw 1420
#27
       tomosynthes*:ti,ab,kw 33
#28
       {or #7-#27}
                      70238
#29
       #6 and #28 Publication Year from 2014 to 2017 772
```

CENTRAL – 694 [RCTs]

HARMS

MEDLINE

Database: Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE(R) Daily and Ovid MEDLINE(R) <1946 to Present> Search Strategy:

- 1 exp Breast Neoplasms/ (287642)
- 2 ((breast* or mamma or mammar*) adj3 (cancer* or carcinoid* or carcinoma* or carcinogen* or adenocarcinoma* or adeno-carcinoma* or malignan* or neoplasia* or neoplasm* or sarcoma* or tumour* or tumor*)).tw,kw. (331761)
- 3 exp Carcinoma, Intraductal, Noninfiltrating/ (10262)
- 4 intraductal carcinoma*.tw,kw. (922)
- 5 (ductal carcinoma in situ or DCIS).tw,kw. (7495)
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- 7 exp Breast Neoplasms/di, pc (47714)
- 8 exp Mass Screening/ (124871)
- 9 screen*.tw,kw. (660022)
- 10 "Early Detection of Cancer"/ (18397)
- 11 ((early or earlier or earliest) adj3 (detect* or diagnos* or identif* or recogni*)).tw,kw.(225345)
- 12 exp Self-Examination/ (2485)

- 13 ((self-exam* or self-detect* or self-screen*) adj5 (breast\$1 or mamma or mammary or nipple\$1)).tw,kw. (2050)
- 14 Physical Examination/ (40906)
- 15 (exam* adj5 (breast? or mamma or mammar* or nipple?)).tw,kw. (15055)
- 16 exp Breast Neoplasms/ra (16756)
- 17 exp Mammography/ (32349)
- 18 (mammograph* or mammogram*).tw,kw. (33182)
- 19 exp Magnetic Resonance Imaging/ (415717)
- 20 (fMRI or fMRIs or MRI or MRIs or NMRI or NMRIs or MR imaging or NMR imaging or magnetic resonance imag* or magnetic resonance tomograph* or MR tomograph*).tw,kw. (380636)
- 21 (chemical shift imaging or proton spin tomograph* or zeugmatograph*).tw,kw. (1076)
- 22 exp Breast Neoplasms/us (4023)
- 23 (ultrasound* or ultrason* or echograph* or echomammogra* or echo-mammogra* or echotomograph* or echo-tomograph* or sonograph*).tw,kw. (382288)
- 24 Imaging, Three-Dimensional/ (64456)
- 25 ((3D or "3-D") adj3 imag*).tw,kw. (17743)
- 26 (("3" or three) adj dimension* adj3 imag*).tw,kw. (15527)
- 27 tomosynthes*.tw,kw. (1236)
- 28 or/7-27 (1875233)
- 29 6 and 28 [BREAST CANCER SCREENING] (102429)
- 30 Male/ not (Female/ and Male/) (2788208)
- 31 29 not 30 [MALE-ONLY REMOVED] (100934)
- 32 exp Infant/ not (exp Adult/ and exp Infant/) (838449)
- 33 exp Child/ not (exp Adult/ and exp Child/) (1197384)
- 34 Adolescent/ not (exp Adult/ and Adolescent/) (595774)
- 35 or/32-34 (1865046)
- 36 31 not 35 [CHILD-ONLY REMOVED] (100454)
- 37 exp Animals/ not (exp Animals/ and Humans/) (4850259)
- 38 36 not 37 [ANIMAL-ONLY REMOVED] (98888)
- 39 (comment or editorial or news or newspaper article).pt. (1254980)
- 40 (letter not (letter and randomized controlled trial)).pt. (1008588)
- 41 38 not (39 or 40) [OPINION PIECES REMOVED] (91963)
- 42 (201410* or 201411* or 201412* or 2015* or 2016* or 2017*).dc. (3219115)
- 43 41 and 42 [UPDATE PERIOD] (13074)
- 44 (controlled clinical trial or randomized controlled trial or pragmatic clinical trial).pt. (600336)
- 45 clinical trials as topic.sh. (197690)
- 46 (randomi#ed or randomly or RCT\$1 or placebo*).tw. (882744)
- 47 ((singl* or doubl* or trebl* or tripl*) adj (mask* or blind* or dumm*)).tw. (167447)
- 48 trial.ti. (201433)
- 49 or/44-48 (1273229)
- 50 43 and 49 [RCTS] (1017)
- 51 remove duplicates from 50 [RCTS DUPLICATES REMOVED] (738)
- 52 controlled clinical trial.pt. (98123)
- 53 Controlled Clinical Trial/ or Controlled Clinical Trials as Topic/ (103798)
- 54 (control* adj2 trial*).tw. (231865)
- 55 Non-Randomized Controlled Trials as Topic/ (135)
- 56 (nonrandom* or non-random* or quasi-random* or quasi-experiment*).tw. (49091)
- 57 (nRCT or nRCTs or non-RCT\$1).tw. (605)

```
58 (pre- adj3 post-).tw. (65615)
59 (pretest adj3 posttest).tw. (4480)
60 Historically Controlled Study/(111)
61
    (control* adj2 stud$3).tw. (212253)
62 Control Groups/ (1833)
63 (control$ adj2 group$1).tw. (436609)
64
    trial.ti. (201433)
    or/52-64 (1085914)
65
66
    43 and 65 [NON-RCTS] (1128)
    66 not 50 [OVERLAP WITH RCT SET REMOVED] (496)
67
68
    remove duplicates from 67 [NON-RCTS - DUPLICATES REMOVED] (415)
69
    exp Cohort Studies/ (1803827)
70
    cohort$1.tw. (470255)
71
    Retrospective Studies/ (674584)
72
    (longitudinal or prospective or retrospective).tw. (1065169)
73
    ((followup or follow-up) adj (study or studies)).tw. (48662)
    Observational study.pt. (35331)
75
    (observation$2 adj (study or studies)).tw. (78822)
76
    ((population or population-based) adj (study or studies or analys#s)).tw. (15420)
    ((multidimensional or multi-dimensional) adj (study or studies)).tw. (96)
77
78 Comparative Study.pt. (1958641)
    ((comparative or comparison) adj (study or studies)).tw. (101181)
79
80
    exp Case-Control Studies/ (917378)
    ((case-control* or case-based or case-comparison) adj (study or studies)).tw. (95907)
81
82
    (ecolog* adj (study or studies)).tw. (4768)
83
    or/69-82 (4242593)
84
    43 and 83 [OBSERVATIONAL STUDIES] (4241)
    84 not (50 or 66) [OVERLAP WITH RCTS AND NON-RCTS REMOVED] (3507)
85
    remove duplicates from 85 [OBSERVATIONAL STUDIES - DUPLICATES REMOVED] (2709)
86
87
    exp Mass Screening/ae [Adverse Effects] (800)
    "Early Detection of Cancer"/ae [Adverse Effects] (247)
88
89
    exp Self-Examination/ae [Adverse Effects] (2)
90
    exp Mammography/ae [Adverse Effects] (805)
    exp Diagnostic Errors/(118284)
91
92
    misdiagnos*.tw,kw. (27228)
93
    (miss$2 adj3 diagnos*).tw,kw. (4750)
94
    (overdiagnos* or over diagnos*).tw,kw. (4548)
95
    (false adj (negative* or positive*)).tw,kw. (73344)
96 ((error* or false$2 or wrong$2) adj3 (alarm* or detect* or diagnos*)).tw,kw. (22187)
97
    exp Medical Overuse/ (5464)
98 overtreat*.tw,kw. (3916)
    ((inappropriate* or unnecessar*) adj3 (followup or follow-up or procedur* or therap* or
treatment*)).tw,kw. (11197)
100 (inappropriate* or unnecessar* or safe or adverse or adversely or undesirabl* or unintend* or
unintent* or unwanted or harm* or injurious* or risk or risks or reaction* or complication*).ti. (844248)
101 ((adverse* or undesirabl* or unintend* or unintent* or unwanted or harm* or toxic or injurious*
or serious* or fatal) adj5 (affect or affected or affecting or affects or consequence* or effect* or react or
reacts or reacted or reacting or reaction* or event* or outcome* or incident*)).tw,kw. (547440)
```

```
((adverse* or inappropriat* or unnecessar* or undesirabl* or unintend* or unintent* or
unwanted or injurious* or serious*) adj5 (alarm* or anxiet* or anxious* or distress* or emotion* or
feeling* or psycholog* or uncertaint*)).tw,kw. (7420)
     iatrogen*.tw,kw. (29231)
     or/87-103 (1569802)
104
105
    43 and 104 [HARMS OF BREAST CANCER SCREENING] (2235)
106
     105 and 51 [HARMS OF BREAST CANCER SCREENING - RCTS] (206)
107
     105 and 68 [HARMS OF BREAST CANCER SCREENING - NON-RCTS] (120)
108
     105 and 86 [HARMS OF BREAST CANCER SCREENING - OBSERVATIONAL STUDIES] (567)
109 or/106-108 [HARMS OF BREAST CANCER SCREENING - ALL STUDY DESIGNS] (893)
Cochrane Library
Search Name: CTFPHC - Breast Cancer Screening - All Modalities - Harms
Date Run:
              04/01/17 17:41:41.934
              2017 Jan 4 (OHRI) - Oct 2014-present - FINAL
Description:
ID
       Search Hits
#1
       [mh "Breast Neoplasms"]
                                     9949
       ((breast* or mamma or mammar*) near/3 (cancer* or carcinoid* or carcinoma* or carcinogen*
#2
or adenocarcinoma* or adeno-carcinoma* or malignan* or neoplasia* or neoplasm* or sarcoma* or
tumour* or tumor*)):ti,ab,kw 22627
#3
       [mh "Carcinoma, Intraductal, Noninfiltrating"]
#4
       (intraductal next carcinoma*):ti,ab,kw 181
#5
       ("ductal carcinoma in situ" or DCIS):ti,ab,kw
                                                    302
#6
       {or #1-#5}
                      22683
#7
       [mh "Breast Neoplasms"/DI,PC] 1459
       [mh "Mass Screening"] 5540
#8
#9
       screen*:ti,ab,kw
                              29461
       [mh "Early Detection of Cancer"]
#10
                                             898
#11
       ((early or earlier or earliest) near/3 (detect* or diagnos* or identif* or recogni*)):ti,ab,kw
       5661
#12
       [mh Self-Examination] 202
#13
       ((self next (exam* or detect* or screen*)) near/5 (breast* or mamma or mammary or
nipple*)):ti,ab,kw
                      208
       [mh ^"Physical Examination"] 913
#14
#15
       (exam* near/5 (breast* or mamma or mammar* or nipple*)) .tw,kw.
                                                                           2
#16
       [mh "Breast Neoplasms"/RA]
                                     380
#17
       [mh Mammography]
                             1033
#18
       (mammograph* or mammogram*):ti,ab,kw
                                                    1859
#19
       [mh "Magnetic Resonance Imaging"]
                                             7076
#20
       (fMRI or fMRIs or MRI or MRIs or NMRI or NMRIs or "MR imaging" or "NMR imaging" or
("magnetic resonance" next imaging) or ("magnetic resonance" next tomograph*) or (MR next
tomograph*)):ti,ab,kw 14833
#21
       ("chemical shift imaging" or ("proton spin" next tomograph*) or zeugmatograph*):ti,ab,kw
       20
       [mh "Breast Neoplasms"/US]
#22
```

```
#23
       (ultrasound* or ultrason* or echograph* or echomammogra* or echo-mammogra* or
echotomograph* or echo-tomograph* or sonograph*):ti,ab,kw 21358
#24
       [mh "Imaging, Three-Dimensional"]
                                              1022
#25
       ((3D or "3-D") near/3 imag*):ti,ab,kw
                                              338
#26
       (((3 or three) next dimension*) near/3 imag*):ti,ab,kw 1420
#27
       tomosynthes*:ti,ab,kw 33
#28
       {or #7-#27}
                       70238
#29
       #6 and #28
                       3794
#30
       [mh "Mass Screening"/AE]
                                       45
#31
       [mh "Early Detection of Cancer"/AE]
                                              13
#32
       [mh Self-Examination/AE]
                                      0
#33
       [mh Mammography/AE]
                                       28
#34
       [mh "Diagnostic Errors"]
                                       2916
#35
       misdiagnos*:ti,ab,kw 210
#36
       (miss* near/3 diagnos*):ti,ab,kw
                                              92
#37
       (overdiagnos* or (over next diagnos*)):ti,ab,kw 190
#38
       (false next (negative* or positive*)):ti,ab,kw
#39
       ((error* or false* or wrong*) near/3 (alarm* or detect* or diagnos*)):ti,ab,kw
                                                                                     1187
#40
       [mh "Medical Overuse"]
                                       138
#41
       overtreat*:ti,ab,kw
                               193
#42
       ((inappropriate* or unnecessar*) near/3 (followup or "follow-up" or procedur* or therap* or
treatment*)):ti,ab,kw 564
#43
       (inappropriate* or unnecessar* or safe or adverse or adversely or undesirabl* or unintend* or
unintent* or unwanted or harm* or injurious* or risk or risks or reaction* or complication*):ti
       ((adverse* or undesirabl* or unintend* or unintent* or unwanted or harm* or toxic or
injurious* or serious* or fatal) near/5 (affect or affected or affecting or affects or consequence* or
effect* or react or reacts or reacted or reacting or reaction* or event* or outcome* or
incident*)):ti,ab,kw
                       122393
       ((adverse* or inappropriat* or unnecessar* or undesirabl* or unintend* or unintent* or
#45
unwanted or injurious* or serious*) near/5 (alarm* or anxiet* or anxious* or distress* or emotion* or
feeling* or psycholog* or uncertaint*)):ti,ab,kw 1201
       iatrogen*:ti,ab,kw
#46
                               691
#47
       {or #30-#46}
                       160469
       #29 and #47 Publication Year from 2014 to 2017229
#48
```

CENTRAL – 216 [RCTs]

BREAST SELF-EXAM – Missed Search Period

MEDLINE

Database: Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE(R) Daily and Ovid MEDLINE(R) <1946 to Present> Search Strategy:

¹ exp Breast Neoplasms/ (287642)

- 2 ((breast* or mamma or mammar*) adj3 (cancer* or carcinoid* or carcinoma* or carcinogen* or adenocarcinoma* or adeno-carcinoma* or malignan* or neoplasia* or neoplasm* or sarcoma* or tumour* or tumor*)).tw,kw. (331761)
- 3 exp Carcinoma, Intraductal, Noninfiltrating/ (10262)
- 4 intraductal carcinoma*.tw,kw. (922)
- 5 (ductal carcinoma in situ or DCIS).tw,kw. (7495)
- 6 or/1-5 [BREAST CANCER] (395424)
- 7 exp Self-Examination/ (2485)
- 8 ((self-exam* or self-detect* or self-screen*) adj5 (breast\$1 or mamma or mammary or nipple\$1)).tw,kw. (2050)
- 9 or/7-8 (3569)
- 10 6 and 9 [BREAST SELF-EXAMINATION] (2287)
- 11 Male/ not (Female/ and Male/) (2788208)
- 12 10 not 11 (2280)
- 13 exp Infant/ not (exp Adult/ and exp Infant/) (838449)
- 14 exp Child/ not (exp Adult/ and exp Child/) (1197384)
- 15 Adolescent/ not (exp Adult/ and Adolescent/) (595774)
- 16 or/13-15 (1865046)
- 17 12 not 16 (2259)
- 18 exp Animals/ not (exp Animals/ and Humans/) (4850259)
- 19 17 not 18 [ANIMAL-ONLY REMOVED] (2259)
- 20 (comment or editorial or news or newspaper article).pt. (1254980)
- 21 (letter not (letter and randomized controlled trial)).pt. (1008588)
- 22 19 not (20 or 21) [OPINION PIECES REMOVED] (2114)
- 23 (201010* or 201011* or 201012* or 2011* or 2012* or 2013* or 201401* or 201402* or 201403* or 201404* or 201405* or 201406* or 201407* or 201408* or 201409*).dc. (4562092)
- 24 22 and 23 [UPDATE PERIOD] (297)
- 25 (controlled clinical trial or randomized controlled trial or pragmatic clinical trial).pt. (600336)
- 26 clinical trials as topic.sh. (197690)
- 27 (randomi#ed or randomly or RCT\$1 or placebo*).tw. (882744)
- 28 ((singl* or doubl* or trebl* or tripl*) adj (mask* or blind* or dumm*)).tw. (167447)
- 29 trial.ti. (201433)
- 30 or/25-29 (1273229)
- 31 24 and 30 [RCTS] (26)
- 32 remove duplicates from 31 [RCTS DUPLICATES REMOVED] (24)
- 33 controlled clinical trial.pt. (98123)
- 34 Controlled Clinical Trial/ or Controlled Clinical Trials as Topic/ (103798)
- 35 (control* adj2 trial*).tw. (231865)
- 36 Non-Randomized Controlled Trials as Topic/ (135)
- 37 (nonrandom* or non-random* or quasi-random* or quasi-experiment*).tw. (49091)
- 38 (nRCT or nRCTs or non-RCT\$1).tw. (605)
- 39 (pre- adj3 post-).tw. (65615)
- 40 (pretest adj3 posttest).tw. (4480)
- 41 Historically Controlled Study/ (111)
- 42 (control* adj2 stud\$3).tw. (212253)
- 43 Control Groups/ (1833)
- 44 (control\$ adj2 group\$1).tw. (436609)
- 45 trial.ti. (201433)

```
46 or/33-45 (1085914)
47 24 and 46 [NON-RCTS] (19)
48 47 not 31 [OVERLAP WITH RCTS REMOVED] (11)
49
    remove duplicates from 48 [NON-RCTS - DUPLICATES REMOVED] (10)
    exp Cohort Studies/ (1803827)
50
51 cohort$1.tw. (470255)
52
    Retrospective Studies/ (674584)
53
    (longitudinal or prospective or retrospective).tw. (1065169)
54
    ((followup or follow-up) adj (study or studies)).tw. (48662)
55
    Observational study.pt. (35331)
56
    (observation$2 adj (study or studies)).tw. (78822)
57
    ((population or population-based) adj (study or studies or analys#s)).tw. (15420)
    ((multidimensional or multi-dimensional) adj (study or studies)).tw. (96)
58
59
    Comparative Study.pt. (1958641)
60
    ((comparative or comparison) adj (study or studies)).tw. (101181)
    exp Case-Control Studies/ (917378)
61
62 ((case-control* or case-based or case-comparison) adj (study or studies)).tw. (95907)
63
    or/50-62 [OBSERVATIONAL STUDIES] (4238905)
    24 and 63 [OBSERVATIONAL STUDIES] (69)
64
    64 not (31 or 47) [OVERLAP WITH RCTS AND NON-RCTS REMOVED] (59)
65
66 remove duplicates from 65 [OBSERVATIONAL STUDIES - DUPLICATES REMOVED] (53)
    32 or 49 or 66 [ALL STUDY DESIGNS] (87)
67
*********
Cochrane Library
Search Name: CTFPHC - Breast Cancer Screening - Self-Examination
              04/01/17 17:44:19.792
Date Run:
Description:
              2017 Jan 4 - 2010-2014 - FINAL
ID
       Search Hits
#1
       [mh "Breast Neoplasms"]
                                     9949
       ((breast* or mamma or mammar*) near/3 (cancer* or carcinoid* or carcinoma* or carcinogen*
#2
or adenocarcinoma* or adeno-carcinoma* or malignan* or neoplasia* or neoplasm* or sarcoma* or
tumour* or tumor*)):ti,ab,kw 22627
#3
       [mh "Carcinoma, Intraductal, Noninfiltrating"]
                                                    118
#4
       (intraductal next carcinoma*):ti,ab,kw 181
#5
       ("ductal carcinoma in situ" or DCIS):ti,ab,kw
                                                    302
#6
                      22683
       {or #1-#5}
#7
       [mh Self-Examination] 202
#8
       ((self next (exam* or detect* or screen*)) near/5 (breast* or mamma or mammary or
nipple*)):ti,ab,kw
                      208
#9
       #7 or #8
                      303
```

CENTRAL - 23 [RCTs]

#6 and #9 Publication Year from 2010 to 2014 23

#10

Appendix 3- Screening Forms (Updated Search)

	1 – Title and abstract screening Does this record focus on breast cancer screening in a population screening context?
	O Yes/possibly O No*
	O Unclear/no abstract
*Reason	ns for selecting 'no':
	not focus on breast cancer screening in a population screening context (If >20% of the population are high risk- then . For now, include all studies which assess dense breasts populations).
ovarian	\underline{k} : women with pre-existing or personal history of breast cancer, family history (in a first degree relative) of breast or cancer or other personal risk factors, such as abnormal breast pathology or BRCA1/BRCA2 genetic mutations, previous d radiation treatment to the chest (such as Hodgkin's) for cancer.
2) Anima	al/in vivo studies
	uses on breast cancer screening but it is clearly obvious that it is one of the following: CPG, SRs, Narrative literature commentary (without primary data), editorials (without primary data), protocol
*Those	e answered yes/unclear will be passed through to full-text screening.
Level 2	2 – Full-text screening
1.	Is the full-text available?
	O Yes
	O No
	O abstract onlyO article not required due to known foreign language
2.	Is the article published in English or French? • Yes
	⊙ No
3.	Is the article any of the following study designs?
	RCTs (including cluster), or novel/extended analysis of RCT data. Non-RCTs
	Comparative cohort studies (including adminstrative database studies/registries) Ecological studies
	Example of studies to exclude:
	case-control, cross-sectional studies,

controlled before-after, diagnostic test accuracy studies modelling studies. Also exclude narrative reviews, systematic reviews/meta-analysis, commentaries & Editorials (without primary data), protocols, papers on study design O Yes O No O Diagnostic Type Accuracy Study- of the interventions themselves, exclude kappa studies on observer agreement 4. Is the article focused on breast cancer screening (must mention inclusion of some sort of screening practice)? Exclude: (i) studies where focus of the intervention is to randomize patients to programs to enforce/enhance screening. Ex: community health worker-led health literacy intervention; (ii) studies on treatment O Yes O No 5. Is it the population of interest? O No- women <40 years (exclusively) O No- women ≥ 40 years who are high –risk (based on family history and other personal risk factors- genetic mutations, abnormal pathology, previous history of cancer, etc). O Yes- women ≥ 40 years who are 'not at high risk'- i.e., average risk (or at least80% of the population is not at high risk) O Yes- women ≥ 40 years who have dense breasts (>75% of population) O Unclear- mixed aged population who are 'not at high risk' (at least 80% of the population) or who have dense breasts (<75% of the population) O No-mixed aged population who are at 'high risk' (>20% of population) or dense breasts (>75%) 6. Does it include the intervention of interest? Mammography (film, digital, tomosynthesis) with or without CBE/BSE MRI with or without CBE/BSE Ultrasound with or without CBE/BSE CBE **BSE** O Yes O No 7. Is the comparator: "no screening", "usual care"?

case-series.

O Yes
O No

Typically, these questions are nested. If an answer allows us to proceed in the inclusion criteria, the next question will appear. Those bolded would be those that would pass through to the following question. If question 7 is 'Yes', this article would be passed through to a post-hoc evaluation, ensuring it has outcomes of interest.

Appendix 4- Data Extraction- Overview of Reviews

Publication details: year of publication, language, publication status

Search details: databases searched and years searched

Selection criteria: Number of included studies, type of study design, population, sample sizes, quality of included studies (must align with the CTFPHC PICOTs)

Results of the systematic review: summarize qualitatively body of evidence

Results of the meta-analysis: pooled estimate, heterogeneity tests

Stregnths of limitations of the review

AMSTAR quality

Appendix 5- Assessing the Methodological Quality of Systematic Reviews (AMSTAR) (Overview of Reviews)

1. Was an 'a priori' design provided? The research question and inclusion criteria should be established before the conduct of the review. Note: Need to refer to a protocol, ethics approval, or predetermined/a priori published research objectives to score a "yes."	ansv	Yes No Can't wer Not licable
2. Was there duplicate study selection and data extraction? There should be at least two independent data extractors and a consensus procedure for disagreements should be in place. Note: 2 people do study selection, 2 people do data extraction, consensus process or one person checks the other's work.	ansv appl	Yes No Can't wer Not licable
3. Was a comprehensive literature search performed? At least two electronic sources should be searched. The report must include years and databases used (e.g., Central, EMBASE, and MEDLINE). Key words and/or MESH terms must be stated and where feasible the search strategy should be provided. All searches should be supplemented by consulting current contents, reviews, textbooks, specialized registers, or experts in the particular field of study, and by reviewing the references in the studies found. Note: If at least 2 sources + one supplementary strategy used, select "yes" (Cochrane register/Central counts as 2 sources; a grey literature search counts as supplementary).	ansv	Yes No Can't wer Not licable
4. Was the status of publication (i.e. grey literature) used as an inclusion criterion? The authors should state that they searched for reports regardless of their publication type. The authors should state whether or not they excluded any reports (from the systematic review), based on their publication status, language etc.	ansv	Yes No Can't wer

Note: If review indicates that there was a search for "grey literature" or "unpublished literature," indicate "yes." SINGLE database, dissertations, conference proceedings, and trial registries are all considered grey for this purpose. If searching a source that contains both grey and non-grey, must specify that they were searching for grey/unpublished lit.	app	Not licable
		Yes
5. Was a list of studies (included and excluded) provided? A list of included and excluded studies should be provided.		No
Note: Acceptable if the excluded studies are referenced. If there is an electronic link to the list but the link is dead, select "no."	ansv app	Can't wer Not licable
6. Were the characteristics of the included studies provided? In an aggregated form such as a table, data from the original		Yes
studies should be provided on the participants, interventions and outcomes. The ranges of characteristics in all the studies analyzed		No
e.g., age, race, sex, relevant socioeconomic data, disease status, duration, severity, or other diseases should be reported.	ansv	Can't wer
Note: Acceptable if not in table format as long as they are described as above.	app	Not licable
7. Was the scientific quality of the included studies assessed and documented? 'A priori' methods of assessment should be provided (e.g., for effectiveness studies if the author(s) chose to include only randomized, double-blind, placebo controlled studies, or allocation concealment as inclusion criteria); for other types of studies alternative items will be relevant.		Yes No
Note: Can include use of a quality scoring tool or checklist, e.g., Jadad scale, risk of bias, sensitivity analysis, etc., or a description of quality items, with some kind of result for EACH study ("low" or "high" is fine, as long as it is clear which studies scored "low" and which scored "high"; a summary score/range for all studies is not acceptable).	ansv app	Can't wer Not licable
8. Was the scientific quality of the included studies used appropriately in formulating conclusions?		Yes
The results of the methodological rigor and scientific quality should		No

be considered in the analysis and the conclusions of the review, and explicitly stated in formulating recommendations. Note: Might say something such as "the results should be interpreted with caution due to poor quality of included studies." Cannot score "yes" for this question if scored "no" for question 7.	ansv ansv app	Can't wer Not licable
9. Were the methods used to combine the findings of studies appropriate? For the pooled results, a test should be done to ensure the studies were combinable, to assess their homogeneity (i.e., Chi-squared test for homogeneity, I2). If heterogeneity exists a random effects model should be used and/or the clinical appropriateness of combining should be taken into consideration (i.e., is it sensible to		Yes No Can't
combine?). Note: Indicate "yes" if they mention or describe heterogeneity, i.e., if they explain that they cannot pool because of heterogeneity/variability between interventions.	ansv	wer Not licable
10. Was the likelihood of publication bias assessed? An assessment of publication bias should include a combination of graphical aids (e.g., funnel plot, other available tests) and/or statistical tests (e.g., Egger regression test, Hedges-Olken). Note: If no test values or funnel plot included, score "no". Score "yes" if mentions that publication bias could not be assessed because there were fewer than 10 included studies.	ansv app	Yes No Can't wer Not licable
11. Was the conflict of interest included? Potential sources of support should be clearly acknowledged in both the systematic review and the included studies. Note: To get a "yes," must indicate source of funding or support for the systematic review AND for each of the included studies.	ansv	Yes No Can't wer Not licable

Shea et al. BMC Medical Research Methodology 2007 7:10 doi:10.1186/1471-2288-7-10

Appendix 6- Cochrane Risk of Bias Tool

1.	Selection bias domain: Random sequence generation O Low risk O Unclear risk O High risk					
	Support for judgement:					
2.	Selection bias domain: Allocation concealment					
	O Low risk					
	O Unclear risk					
	O High risk					
	Support for judgement:					
3.	Performance bias domain: Blinding of participants and personnel (for each outcome)					
	O Low risk					
	O Unclear risk					
	O High risk					
	Support for judgement:					
4.	Detection bias domain: Blinding of outcome assessment (for each outcome)					
	O Low risk					
	O Unclear risk					
	O High risk					
	Support for judgement:					

5.	Attrition bias domain: Incomplete outcome data (for each outcome)
	O Low risk
	O Unclear risk
	O High risk
	Support for judgement:
6.	Reporting bias domain: Selective reporting
	O Low risk
	O Unclear risk
	O High risk
	Support for judgement:
7.	Other sources of bias
•	O Low risk
	O Unclear risk
	O High risk
	Support for judgement:
	Support for judgement.

Appendix 7- Newcastle-Ottawa Scale (Cohort Studies)

<u>Note</u>: A study can be awarded a maximum of one star for each numbered item within the Selection and Outcome categories. A maximum of two stars can be given for Comparability

Selection	
1) Representativeness of the exposed cohort	
a) truly representative of the average	(describe) in the community ★
b) somewhat representative of the average	
c) selected group of users eg nurses, volunteers	
d) no description of the derivation of the cohort	
2) <u>Selection of the non-exposed cohort</u>	
a) drawn from the same community as the exposed co	ohort *
b) drawn from a different source	
c) no description of the derivation of the non-exposed	cohort
3) <u>Ascertainment of exposure</u>	
a) secure record (eg surgical records) ★	
b) structured interview ★	
c) written self-report	
d) no description	
4) Demonstration that outcome of interest was not present at s	start of study
a) yes ≭	
b) no	
Comparability	
1) Comparability of cohorts on the basis of the design or analysi	<u>iis</u>
a) study controls for(select the most in	
b) study controls for any additional factor $lpha$ (This crite	eria could be modified to indicate specific control
for a second important factor.)	
* Age and Hormone replacement therapy use w	were considered.
Outcome	
1) Assessment of outcome	
a) independent blind assessment *	
b) record linkage ∗	
c) self report	
d) no description	
2) Was follow-up long enough for outcomes to occur	
a) yes (select an adequate follow up period for outcom	ne of interest) *
b) no	
3) Adequacy of follow up of cohorts	
a) complete follow up - all subjects accounted for $igspace{*}$	
b) subjects lost to follow up unlikely to introduce bias -	
adequate %) follow up, or description provided of thos	se lost) ★
c) follow up rate <% (select an adequate %) and n	no description of those lost
d) no statement	

- *Modified tool to add one more question under 'outcome'
 4) Did the authors adjust for lead time bias in the analysis (or was follow-up long-enough to reduce lead time bias)?
 - a) yes*
 - b) no

Appendix 9 – List of Excluded Studies (Full Text) (Updated Search)

Full Text Unavailable

RefID:2874. Boonyaleepan, Araya. Positron Emission Mammography for Breast Cancer in Rajavithi Hospital. Journal of the Medical Association of Thailand = Chotmaihet thangphaet 2016. 99 Suppl 2 () S130-S135. Full Text Unavailable

RefID:984. Luijt, P., Heijnsdijk, E. A. M., Fracheboud, J., Broeders, M. J. M., Wesseling, J., Heeten, G. J., and Koning, H. J.. DCIS distribution of grades in 5,126 screened and non-screened women and estimated risk of overdiagnosis in breast cancer screening: A model of progression. European journal of cancerConference Abstract 2014. 50 () S168-.

RefID:2703. Menes, Tehillah S., Kerlikowske, Karla, Lange, Jane, Jaffer, Shabnam, Rosenberg, Robert, and Miglioretti, Diana L.. Subsequent Breast Cancer Risk Following Diagnosis of Atypical Ductal Hyperplasia on Needle Biopsy. JAMA oncology 2017. 3 (1) 36-41.

RefID:1311. Simmons, R.. Long-term results of phase II ablation after breast lumpectomy added to extend intraoperative margins (ABLATE I) trial. Breast DiseasesNote 2015. 25 (4) 331-332.

Abstract Only

RefID:844. Autier, P., Boniol, M., Smans, M., and Boyle, P.. Randomized trials on mammography screening and the left-to-nature design. Journal of clinical oncologyConference Abstract 2014. 32 (15 Suppl 1) -.

RefID:212. Barrajon, E., Lopez, A., and Adrover, E.. Screening mammography in old women saves lives: A simulation model. Journal of clinical oncology: official journal of the American Society of Clinical Oncology 2006. 24 (18_suppl) 10561-.

RefID:101. Bonanni, B., Maisonneuve, P., Serrano, D., Varricchio, C., Cazzaniga, M., Lazzeroni, M., Santillo, B., Di Pace, R., Meneghetti, L., Tagliafico, A., Veronesi, U., and De Censi, A.. Safety and efficacy of HRT and low-dose tamoxifen in a phase II trial (HOT): Analysis of mammographic density and endometrial thickness. Journal of clinical oncology: official journal of the American Society of Clinical Oncology 2011. 29 (15_suppl) 1527-.

RefID:1191. Chan, E. K., Wilson, C., Tyldesley, S., Lai, A., Sam, J., Harry, R., and Nichol, A.. Improving screening mammography return rates in overdue women: A randomized study of signed reminder letters from family physicians. Journal of clinical oncologyConference Abstract 2014. 32 (26 Suppl 1) -.

RefID:3056. Chung, Alice, Gangi, Alexandra, Amersi, Farin, Zhang, Xiao, and Giuliano, Armando. Not Performing a Sentinel Node Biopsy for Older Patients With Early-Stage Invasive Breast Cancer. JAMA surgery 2015. 150 (7) 683-684

RefID:1197. Cyr, A., Tucker, N., Gao, F., Margenthaler, J., Aft, R., Eberlein, T., Appleton, C., Reichert, V., and Gillanders, W.. Pilot phase study results of a prospective, randomized controlled clinical trial evaluating axillary ultrasound vs sentinel lymph node biopsy for axillary staging in early-stage breast cancer patients. Annals of surgical oncologyConference Abstract 2015. 22 (2 Suppl 1) 14-15.

RefID:2753. Dawson, S., McKinley, J., Jenkins, M., McLachlan, S., Lindeman, G., Friedlander, M., Hopper, J., and Phillips, K.. Cancer risk management practices of non-carriers within BRCA1/2 mutation positive families in the Kathleen Cunningham Consortium for Research into Familial Breast Cancer (kConFab). Journal of clinical oncology: official journal of the American Society of Clinical Oncology 2006. 24 (18_suppl) 1020-.

RefID:1217. Diaz-Santana, M. V. and Reeves, K. W.. Breast cancer risk factors and screening practices among Hispanics subgroups in the United States. Cancer Epidemiology Biomarkers and Prevention. Conference: 7th AACR Conference on the Science of Health Disparities in Racial/Ethnic Minorities and the Medically Underserved San Antonio, TX United States. Conference Start: 20141109 Conference End: 2014111Conference Abstract 2015. 24 (10 Suppl 1 no pagination) -.

RefID:976. Durham, D., Robinson, W., Lee, S., Wheeler, S., Bowling, J., and Henderson, L.. Disparities in time to diagnostic follow up after screening mammography. Cancer Epidemiology Biomarkers and Prevention. Conference: 7th AACR Conference on the Science of Health Disparities in Racial/Ethnic Minorities and the Medically Underserved San Antonio, TX United States. Conference Start: 20141109 Conference End: 2014111Conference Abstract 2015. 24 (10 Suppl 1 no pagination) -.

- RefID:935. Elshof, L. E., Tryfonidis, K., Slaets, L., Leeuwen-Stok, A. E., Dif, N., Skinner, V. P., Loo, C. E., Warnars, G., Bleiker, E., Pijnappel, R. M., Bijker, N., Rutgers, E. J. T., and Wesseling, J.. The LORD trial: A randomized, non-inferiority trial, between active surveillance versus standard treatment in patients with low risk ductal carcinoma in situ. Cancer researchConference Abstract 2015. 75 (9 Suppl 1) -.
- RefID:1071. Henderson, L. M., Benefield, T., Marsh, M. W., and Nakayoshi, M.. Performance of digital diagnostic mammography by race. Cancer Epidemiology Biomarkers and Prevention.Conference: 7th AACR Conference on the Science of Health Disparities in Racial/Ethnic Minorities and the Medically Underserved San Antonio, TX United States.Conference Start: 20141109 Conference End: 2014111Conference Abstract 2015. 24 (10 Suppl 1 no pagination) -.
- RefID:1085. Jones, B. A., Epstein, L., Genao, I., Nunez-Smith, M., Vila, H. S., Claus, E., and Nappi, S.. Perceived control over health and history of mammography screening in Hispanic/Latino women living in the Northeast United States. Cancer Epidemiology Biomarkers and Prevention.Conference: 7th AACR Conference on the Science of Health Disparities in Racial/Ethnic Minorities and the Medically Underserved San Antonio, TX United States.Conference Start: 20141109 Conference End: 2014111Conference Abstract 2015. 24 (10 Suppl 1 no pagination) -.
- RefID:2718. Kim, H., Han, W., Moon, H., Ahn, S. K., Yom, C. K., Shin, H., and Noh, D.. The comparison of the evaluation of axillary lymph node metastasis in breast cancer among PET, chest CT, and ultrasound sonography. Journal of clinical oncology: official journal of the American Society of Clinical Oncology 2011. 29 (15_suppl) e11567-.
- RefID:198. Kirstein, L. J., Keto, J. L., Sanchez, D. P., Fulop, T., Cohen, I., Cohen, J. M., Harshan, M., and Boolbol, S. K.. MRI versus breast-specific gamma imaging (BSGI) in the detection of synchronous breast cancer: A prospective head-to-head trial. Journal of clinical oncology: official journal of the American Society of Clinical Oncology 2011. 29 (27 suppl) 72-.
- RefID:2726. Kojima, S., Hara, A., Kosaka, N., Matsuo, Y., Suzuki, H., Torigoe, S., Suzuki, T., Teramukai, S., Uno, K., and Fukushima, M.. Cancer screening using whole-body 18FDG-PET scan in healthy voluntary subjects. Journal of clinical oncology: official journal of the American Society of Clinical Oncology 2004. 22 (14_suppl) 6072-.
- RefID:1221. Laakmann, E., Witzel, I., Fehm, T., Hesse, T., Minckwitz, G., Mobus, V., Park-Simon, T.-W., Neunhoffer, T., Schmidt, M., Loibl, S., and Muller, V.. Brain metastases in breast cancer network Germany (BMBC, GBG 79): The introduction of the multicenter register and analysis of patient data. Oncology Research and Treatment.Conference: 32.Deutscher Krebskongress, DKK 2016 Berlin Germany.Conference Start: 20160224 Conference End: 20160227.Conference Publication: (var.pagings)Conference Abstract 2016. 39 () 50-.
- RefID:1073. Lee, H. Y., Le, C., Ghebre, R., and Yee, D.. Mobile phone multimedia messaging intervention for breast cancer screening. Cancer researchConference Abstract 2016. 76 (4 Suppl 1 no pagination) -.
- RefID:177. Lin, C., Moore, D., DeMichele, A., Ollila, D., Montgomery, L., Liu, M., Krontiras, H., Gomez, R., Esserman, L., and SPY, TRIAL, I. Detection of locally advanced breast cancer in the I-SPY TRIAL (CALGB 150007/150012, ACRIN 6657) in the interval between routine screening. Journal of clinical oncology: official journal of the American Society of Clinical Oncology 2009. 27 (15_suppl) 1503-.
- RefID:2712. Lowry, H., Dekhne, N., Fend, D., Lerman, R., Gregory, N., and Boura, J.. Multidisciplinary high-risk program: A community hospital's experience. Journal of clinical oncology: official journal of the American Society of Clinical Oncology 2011. 29 (15_suppl) 1562-.
- RefID:176. Mullai, N., Murugesan, N., Burton, L., Goodin, V., and Stout, A.. Risk of noncompliance due to patient discomfort during screening mammogram. Journal of clinical oncology: official journal of the American Society of Clinical Oncology 2009. 27 (15_suppl) 1522-.
- RefID:1113. Narasimmaraj, P. R., Stover, Fiscalini A., Kaplan, C. P., Van't Veer, L. J., Hallada, A. M., Thompson, C. K., Theiner, S., Borowsky, A., Naeim, A., Anton-Culver, H., Lacroix, A., and Esserman, L. J.. A pilot feasibility study of the WISDOM study, a preference-tolerant randomized controlled trial evaluating a risk-based breast cancer screening strategy. Cancer researchConference Abstract 2016. 76 (4 Suppl 1 no pagination) -.
- RefID:1150. Nguyen, K. H., Karliner, L., and Pasick, R.. Disparities in follow up after abnormal mammogram for multiple Asian subpopulations. Cancer Epidemiology Biomarkers and Prevention. Conference: 7th AACR Conference on the Science of Health Disparities in Racial/Ethnic Minorities and the Medically Underserved San Antonio, TX United States. Conference Start: 20141109 Conference End: 2014111Conference Abstract 2015. 24 (10 Suppl 1 no pagination) -.

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Appendix 10- Data Extraction Table (Updated Search)

Study Characteristics				; 2014 for all other modalities)	
*Three publications identified in update, in which some address more than one RCT.					
Breast-Cancer Mortality					
<u> </u>	sual Care/No Screenin				T
1) Trial Name;	1) Age at entry;	1) Type of	1) Definition of	Results	Comments
2) Author;	2) Follow-up for	Mammography; 2)	Outcome; 2)		1) Comments about
3) Study Design; 4)	analysis	# of views; 3) # of	Short-case or		comparator;
Years of study; 5)	3) Total	readers; 4)	Long-case		2)Notes to
End of follow-up; 6)	Randomized	screening interval;	accrual		consider; 3) Other
Country	4) Total	5) # of screening			mortality outcomes
	Randomized	rounds; 6) # of			not relevant for our
	(Intervention,	screens attended,			purposes
	Control)	7) Attendance rate			
1) Malmo I;	1) 45-70;	1) Film;	1) Breast Cancer	45-54:	1) Usual Care & No
2) Nystrom 2016;	2) Mean: 30 years;	2) Two. Starting at	as underlying	# of deaths: INT: 70/8673; CONT: 72/8311; RR:	Screening
3) RCT;	3) 42,283;	round 3, single or	cause of death	0.94 (0.66-1.3)	combination:
4) 1976-NR;	4) INT: 21,088;	two view according	according to the	50-59:	women in the
5) NR;	CONT: 21,195	to parenchymal	Swedish Cause	# of deaths: INT: 76/9285; CONT: 80/9322; RR:	control group, born
6) Sweden		pattern;	of Death	1 (0.73-1.4)	1908-1922, were
		3) 2 readers;	Registry;	55-64:	never invited to
		4) 18-24 months;	2) Short-Case	# of deaths: INT: 53/8194; CONT: 66/8679; RR:	screening, while
		5) Born 1908-1917:	Accrual	0.94 (0.62-1.4)	women born 1923-
		6; Born 1918: 7;		60-70:	1932 were first
		Born 1919-1932: 8; 6) NR;		# of deaths: INT: 32/7816; CONT: 48/7806; RR: 0.73 (0.44-1.2)	invited to screening in 1992-19930-
		7) 73% (1 st round:		45-70 (adjusted for age):	although since SC
		74%)		# of deaths: INT: 130/21088; CONT: 147/21195;	accrual equivalent
				RR: 0.88 (0.70-1.1)	to no screening.;
				45-70 (adjusted for age & including BC deaths	2) N/A;
				not in registry):	3) Weighted
				# of deaths: INT: 146/21088; CONT: 157/21195;	cumulative BC
				RR: 0.93 (0.74-1.2)	mortality per
					100,00 women;
					Numbers needed to
					invite to screen to

					prevent a BC death.
1) Malmo II;	1) 43-49;	1) Film;	1) Breast Cancer	43-49 (adjusted for age):	1) Usual Care- The
2) Nystrom 2016;	2) Mean: 22 years;	2) Two;	as underlying	# of deaths: INT: 38/NR; CONT: 38/NR; RR: 0.85	first screening
3) RCT;	3) 17,793;	3) 2 readers;	cause of death	(0.54-1.3)	round of the control
4) 1978-NR;	4) INT: 9,581;	4) 18-24 months;	according to the	43-49 (adjusted for age & including BC deaths	group took place
5) NR;	CONT: 8,212	5) 1-7	Swedish Cause	not in registry):	between 1991 and
6) Sweden		6) NR;	of Death	# of deaths: INT: 40/NR; CONT: 38/NR; RR: 0.89	1994-although
		7) 73% (1 st round:	Registry;	(0.57-1.4)	since SC accrual
		75-80%)	2) Short-Case		equivalent to no
			Accrual		screening.
					2) N/A;
					3) Weighted
					cumulative BC
					mortality per
					100,00 women;
					Numbers needed to
					invite to screen to
					prevent a BC death.
1) Stockholm;	1) 39-65;	1) NR;	1) Breast Cancer	40-49:	1) Usual Care-
2) Nystrom 2016;	2) Mean: 25 years;	2) One;	as underlying	# of deaths: INT: 29/14303; CONT: 11/8021; RR:	Between 2 nd and 3 rd
3) Quasi- RCT;	3) 60,117;	3) 1 readers;	cause of death	1.5(0.76-3.0)	round between
4) 1981-NR;	4) INT: 39,139;	4) 28 months;	according to the	45-54:	1985 and 1986,
5) NR; 6) Sweden	CONT: 20,978	5) 2 6) NR;	Swedish Cause of Death	# of deaths: INT: 23/14088; CONT: 14/7409; RR: 0.88(0.45-1.7)	control invited to scree-although
		7) 81% (1 st round:	Registry;	50-59:	since SC accrual
		82%)	2) Short-Case	# of deaths: INT: 30/15946; CONT: 26/8421; RR:	equivalent to no
			Accrual	0.61(0.36-1.03)	screening.
				55-65:	2) N/A;
				# of deaths: INT: 47/17357; CONT: 27/8990; RR:	3) Weighted
				0.91(0.56-1.5)	cumulative BC
				40-65 (adjusted for age):	mortality per
				# of deaths: INT: 84/NR; CONT: 48/NR; RR:	100,00 women;
				0.94(0.66-1.3)	Numbers needed to
				40-65 (adjusted for age & including BC deaths	invite to screen to
				not in registry):	prevent a BC death.
				# of deaths: INT: 95/NR; CONT: 56/NR; RR:	
				0.91(0.66-1.3)	
Gothenburg;	1) 39-59;	1) Film;	1) Breast Cancer	40-49:	1)Usual Care-

2) Nystrom 2016; 3) Quasi-RCT; 4) 1982-NR; 5) NR; 6) Sweden	2) Mean: 24 years; 3) 50,200; 4) INT: 21,000; CONT: 29,200	2) 1st round: two & 2nd round: one-two (depending on breast density); 3) 1st-3rd round: 1 reader; 4th-5th rounds: 2 readers; 4) 18 months; 5) Born 1923-1932: 4; Born 1933-1944: 5 6) NR; 7) 81% (1st round: 84%)	as underlying cause of death according to the Swedish Cause of Death Registry; 2) Short-Case Accrual	# of deaths: INT: 30/10888; CONT: 62/13203; RR: 0.59 (0.38-0.90) 45-54: # of deaths: INT: 37/10039; CONT: 65/13518; RR: 0.76 (0.50-1.2) 50-59: # of deaths: INT: 45/10112; CONT: 80/15997; RR: 0.89 (0.60-1.3) 40-59 (adjusted for age): # of deaths: INT: 75/21000; CONT: 142/29200; RR: 0.74 (0.56-0.98) 40-59 (adjusted for age & including BC deaths not in registry): # of deaths: INT: 77/21000; CONT: 149/29200; RR: 0.73 (0.55-0.96)	women in the control group, born from 1923 to 1932 were invited to their first screening round between 1987 and April 1988, and women in the control group born between 1933 and 1944 were invited to their first screening round between February and April 1990-although since SC accrual equivalent to no screening; 2) N/A; 3) Weighted cumulative BC mortality per 100,00 women; Numbers needed to invite to screen to prevent a BC death.
1) UK Age; 2) Moss 2015;	1) 39-41 (invited for screening after 40);	1) NR; 2) 1 st round: two &	1) Defined as deaths with	40+ (Short-case): # of deaths: INT: 182/53883; CONT:	1) Usual Care- invited for
3) RCT;	2) Median: 17.7	Subsequent rounds:	breast cancer	412/106953; RR: 0.88 (0.74-1.04);	screening at age 50
4) 1991-2006;	years (IQR: 16.8-	one (unless	coded as the	Absolute Risk reduction per 1000 women: 0.47	yrs;
5) Dec 31, 2011;	18.8);	otherwise	underlying	(-0.14 to 1.09)	2) N/A;
6) UK	3) 160,921;	indicated);	cause of death	40+ (0-10 years after randomization) (Short-	3) Cumulative BC
	4) INT: 53,914;	3) NR;	on the death	case): # of deaths: INT: 83/53883; CONT: 219/106953;	mortality over time.
	CONT: 107,007	4) 12 months (During NHSBSP-	certificate; 2) Short case	RR: 0.75 (0.59-0.97);	
		every 3 years);	and Long-case	Absolute Risk reduction per 1000 women: 0.51	
		5) NR	accrual	(0.08 to 0.94)	
		6) Mean # of	(separate	40+ (10+ years after randomization) (Short-	

 	1		1
screens attended:	estimates	case):	
4.8 (SD: 3.3)	provided)	# of deaths: INT: 99/53883; CONT: 193/106953;	
7) 81% (at least 1		RR: 1.02 (0.80-1.30);	
routine screen)		Absolute Risk reduction per 1000 women:	
		-0.03 (-0.47 to 0.41)	
		40+ (0-4 years after randomization) (Long-	
		case):	
		# of deaths: INT: 27/NR; CONT: 69/NR; RR: 0.78	
		(0.50-1.21);	
		Absolute Risk reduction per 1000 women:	
		0.14 (-0.10 to 0.39)	
		Absolute Risk reduction per 1000 women years:	
		0.03 (-0.02 to 0.08)	
		40+ (5-9 years after randomization) (Long-	
		case):	
		# of deaths: INT: 56/NR; CONT: 152/NR; RR:	
		0.73 (0.54-0.99);	
		Absolute Risk reduction per 1000 women:	
		0.38 (0.03 to 0.74)	
		Absolute Risk reduction per 1000 women years:	
		0.08 (0.006 to 0.15)	
		40+ (15+ years after randomization) (Long-	
		case):	
		# of deaths: INT: 61/NR; CONT: 109/NR; RR:	
		1.11 (0.81-1.52);	
		Absolute Risk reduction per 1000 women:	
		-0.12 (-0.47 to 0.24)	
		Absolute Risk reduction per 1000 women years:	
		-0.04 (-0.17 to 0.08)	
		40+ (0-17 years after randomization) (Long-	
		case):	
		# of deaths: INT: 242/NR; CONT: 515/NR; RR:	
		0.93 (0.80-1.09);	
		Absolute Risk reduction per 1000 women:	
		0.32 (-0.38 to 1.02)	
		Absolute Risk reduction per 1000 women years:	
		0.02 (-0.02 to 0.06)	

All-Cause Mortality						
Mammography vs. U	Mammography vs. Usual Care/No Screening					
1) UK Age;	1) 39-41 (invited for	1) NR;	1) Defined as	40+:	1)Usual Care-	
2) Moss 2015; 3) RCT; 4) 1991-2006; 5) Dec 31, 2011; 6) UK	screening after 40); 2) Median: 17.7 years (IQR: 16.8- 18.8); 3) 160,921; 4) INT: 53,914; CONT: 107,007	2) 1st round: two & Subsequent rounds: one (unless otherwise indicated); 3) NR; 4) 12 months (During NHSBSPevery 3 years); 5) NR 6) Mean # of screens attended: 4.8 (SD: 3.3) 7) 81% (at least 1 routine screen)	deaths with breast cancer coded as the underlying cause of death on the death certificate (NOTE: nothing mentioned about all-cause); 2) Long-Case Accrual	# of deaths: INT: 2127/53883; CONT: 4320/106953; RR: 0.98 (0.93-1.03)	invited for screening at age 50 yrs; 2) N/A; 3) N/A	
Overdiagnosis		Toutine sereetly				
Mammography + CB	E vs. Usual Care					
1) CNBSS 1& 2 2) Baines 2016; 3) RCT; 4) 1980-1988; 5) Dec 31, 2005; 6) Canada	1) 40-59 (CNBSS 1: 40-49; CNBSS2: 50- 59); 2) Longest follow- up: 25 years; 3) 89,835 (CNBSS1: 50,430; CNBSS2: 39,405); 4) NR	1) NR; 2) NR; 3) NR; 4) 12 months; 5) NR 6) NR 7) CNBSS 1&2- 1st screen: 100% CNBSS1: subsequent screens: INT: 89-86%; CONT: 95-93% CNBSS2: subsequent screens: INT: 90-87%; CONT: 89-85%	1) The numerator is the difference in cancers in the mammography arm compared to the control arm; and the denominator is the # screendetected cancers in the mammography arm; 2) Short case and Long-case accrual (separate estimates	40-49 (Invasive cancer only) (Short-Case): Cum. # of cancer detected: INT: 284; CONT: 225; Difference: 59; Denominator: 213 Estimated Overdiagnosis: 28% 40-49- 1 yr post screening (Invasive cancer only) (Long-Case): Cum. # of cancer detected: INT: 327; CONT: 262; Difference: 65; Denominator: 213 Estimated Overdiagnosis: 31% 40-49- 2 yr post screening (Invasive cancer only) (Long-Case): Cum. # of cancer detected: INT: 379; CONT: 308; Difference: 71; Denominator: 213 Estimated Overdiagnosis: 33% 40-49- 3 yr post screening (Invasive cancer	1) CNBSS1- CBE/Usual Care (single CBE followed by usual care. This constituted a comparison of screening to virtually no screening); CBNSS2- CBE alone; 2) Revised estimates from Miller 2014. Previous publication was confounded by subsequent screening in the	

provided)	only) (Long-Case):	population after
	Cum. # of cancer detected: INT: 435; CONT:	screening ceased in
	363;	1988. They re-
	Difference: 72; Denominator: 213	evaluated the data
	Estimated Overdiagnosis: 34%	by age to provide
	40-49- 4 yr post screening (Invasive cancer	estimates of
	only) (Long-Case):	overdiagnosis at
	Cum. # of cancer detected: INT: 487; CONT:	different time
	421;	points after
	Difference: 66; Denominator: 213	completing of
	Estimated Overdiagnosis: 31%	screening schedules
	40-49- 5 yr post screening (Invasive cancer	in the trial and
	only) (Long-Case):	related them to the
	Cum. # of cancer detected: INT: 544; CONT:	dates at which
	476;	provincial screening
	Difference: 68; Denominator: 213	programs started;
	Estimated Overdiagnosis: 32%	3) N/A
	40-49- 10 yr post screening (Invasive cancer	' '
	only) (Long-Case):	
	Cum. # of cancer detected: INT: 912; CONT:	
	817;	
	Difference: 95; Denominator: 213	
	Estimated Overdiagnosis: 45%	
	40-49- 15 yr post screening (Invasive cancer	
	only) (Long-Case):	
	Cum. # of cancer detected: INT: 1386; CONT:	
	1311;	
	Difference: 75; Denominator: 213	
	Estimated Overdiagnosis: 35%	
	40-49- 15 yr post screening (Invasive cancer	
	only) (Long-Case):	
	Cum. # of cancer detected: INT: 1725; CONT:	
	1622;	
	Difference: 103; Denominator: 213	
	Estimated Overdiagnosis: 48%	
	40-49 (Invasive cancer & In situ) (Short-Case):	
	Cum. # of cancer detected: INT: 326; CONT:	
	234;	

Difference: 92; Denominator: 249 Estimated Overdiagnosis: 37% 40-49- 1 yr post screening (Invasive cancer & In situ) (Long-Case): Cum. # of cancer detected: INT: 371; CONT: Difference: 100; Denominator: 249 Estimated Overdiagnosis: 40% 40-49- 2 yr post screening (Invasive cancer & In situ) (Long-Case): Cum. # of cancer detected: INT: 424; CONT: 318; Difference: 106; Denominator: 249 Estimated Overdiagnosis: 43% 40-49-3 yr post screening (Invasive cancer & In situ) (Long-Case): Cum. # of cancer detected: INT: 480; CONT: Difference: 107; Denominator: 249 Estimated Overdiagnosis: 43% 40-49- 4 yr post screening (Invasive cancer & In situ) (Long-Case): Cum. # of cancer detected: INT: 533; CONT: 432; Difference: 101; Denominator: 249 Estimated Overdiagnosis: 41% 40-49- 5 yr post screening (Invasive cancer & In situ) (Long-Case): Cum. # of cancer detected: INT: 590; CONT: Difference: 103; Denominator: 249 Estimated Overdiagnosis: 41% 40-49- 10 yr post screening (Invasive cancer & In situ) (Long-Case): Cum. # of cancer detected: INT: 958; CONT: 828; Difference: 130; Denominator: 249 Estimated Overdiagnosis: 52%

40-49- 15 yr post screening (Invasive cancer & In situ) (Long-Case): Cum. # of cancer detected: INT: 1432; CONT: 1322; Difference: 110; Denominator: 249 Estimated Overdiagnosis: 44% 40-49- 20 yr post screening (Invasive cancer & In situ) (Long-Case): Cum. # of cancer detected: INT: 1771; CONT: 1633; Difference: 138; Denominator: 249 Estimated Overdiagnosis: 55%
50-59 (Invasive cancer only) (Short-Case): Cum. # of cancer detected: INT: 335; CONT: 256; Difference: 79; Denominator: 271 Estimated Overdiagnosis: 29% 50-59-1 yr post screening (Invasive cancer only) (Long-Case): Cum. # of cancer detected: INT: 381; CONT: 297; Difference: 84; Denominator: 271 Estimated Overdiagnosis: 31% 50-59-2 yr post screening (Invasive cancer only) (Long-Case): Cum. # of cancer detected: INT: 411; CONT: 342; Difference: 69; Denominator: 271 Estimated Overdiagnosis: 25% 50-59-3 yr post screening (Invasive cancer only) (Long-Case): Cum. # of cancer detected: INT: 456; CONT: 399; Difference: 57; Denominator: 271 Estimated Overdiagnosis: 21% 50-59-4 yr post screening (Invasive cancer

cyly (Long Cocc)
only) (Long-Case):
Cum. # of cancer detected: INT: 514; CONT:
468;
Difference: 46; Denominator: 282
Estimated Overdiagnosis: 17%
50-59- 5 yr post screening (Invasive cancer
only) (Long-Case):
Cum. # of cancer detected: INT: 572; CONT:
529;
Difference: 43; Denominator: 271
Estimated Overdiagnosis: 16%
50-59- 10 yr post screening (Invasive cancer
only) (Long-Case):
Cum. # of cancer detected: INT: 899; CONT:
891;
Difference: 8; Denominator: 271
Estimated Overdiagnosis: 3%
50-59- 15 yr post screening (Invasive cancer
only) (Long-Case):
Cum. # of cancer detected: INT: 1295; CONT:
1286;
Difference: 9; Denominator: 271
Estimated Overdiagnosis: 3%
50-59- 15 yr post screening (Invasive cancer
only) (Long-Case):
Cum. # of cancer detected: INT: 1525; CONT:
1511;
Difference: 14; Denominator: 271
Estimated Overdiagnosis: 5%
50-59 (Invasive cancer & In situ) (Short-Case):
Cum. # of cancer detected: INT: 377; CONT:
262;
Difference: 115; Denominator: 312
Estimated Overdiagnosis: 37%
50-59- 1 yr post screening (Invasive cancer & In
situ) (Long-Case):
Cum. # of cancer detected: INT: 424; CONT:

304;
Difference: 120; Denominator: 312
Estimated Overdiagnosis: 38%
50-59- 2 yr post screening (Invasive cancer & In
situ) (Long-Case):
Cum. # of cancer detected: INT: 454; CONT:
349;
Difference: 105; Denominator: 312
Estimated Overdiagnosis: 34%
50-59- 3 yr post screening (Invasive cancer & In
situ) (Long-Case):
Cum. # of cancer detected: INT: 499; CONT:
406;
Difference: 93; Denominator: 312
Estimated Overdiagnosis: 30%
50-59- 4 yr post screening (Invasive cancer & In
situ) (Long-Case):
Cum. # of cancer detected: INT: 557; CONT:
475;
Difference: 82; Denominator: 312
Estimated Overdiagnosis: 26%
50-59- 5 yr post screening (Invasive cancer & In
situ) (Long-Case):
Cum. # of cancer detected: INT: 615; CONT:
536;
Difference: 79; Denominator: 312
Estimated Overdiagnosis: 25%
50-59- 10 yr post screening (Invasive cancer &
In situ) (Long-Case):
Cum. # of cancer detected: INT: 942; CONT:
898;
Difference: 44; Denominator: 312
Estimated Overdiagnosis: 14%
50-59- 15 yr post screening (Invasive cancer &
In situ) (Long-Case):
Cum. # of cancer detected: INT: 1338; CONT:
1293;
Difference: 45; Denominator: 312

Estimated Overdiagnosis: 14% 50-59- 20 yr post screening (Invasive cancer & In situ) (Long-Case):
Cum. # of cancer detected: INT: 1568; CONT:
1518;
Difference: 50; Denominator: 312
Estimated Overdiagnosis: 16%

Appendix 11- Mammography +/- Clincial Breast Exam for Breast-Cancer Mortality (Short-Case Accrual) Forest Plots for Sub-Group Analyses

EVIDENCE SET 1b
Part A- Forest Plot – Breast Cancer Mortality (Short-Case Accrual) (Stratified by CBE use)

Reference	Study	Age (at entry)	Mean Follow-up (yrs)	log [RR]	SE	Weight	Risk Ratio [95%CI]	Risk Ratio (RR) IV, Random, 95% CI
(+) CBE								
Shapiro 1988	HIP	40-64	18.0	-0.24	0.12	12.9%	0.79 [0.63, 1.00]	
Miller 2014	CNBSS 1 & 2	40-59	21.9	0.05	0.11	15.3%	1.05 [0.85, 1.30]	
Subtotal (95%	CI)					28.1%	0.91 [0.69, 1.21]	
Heterogeneity:	$Tau^2 = 0.03$; $Chi^2 = 3.1$	11, df = 1 (F	$P = 0.08$; $I^2 = 68\%$					
Test for overall	effect: Z = 0.63 (P = 0	.53)						
(-)CBE								
Tabar 2011	Swedish Two County	40-74	29.0a	-0.31	0.10	16.2%	0.73 [0.59, 0.90]**	
Nystrom 2016	Gothenburg	40-59	24.0	-0.30	0.14	9.3%	0.74 [0.56, 0.98]	
Nystrom 2016	Malmo II	43-49	22.0	-0.16	0.22	4.0%	0.85 [0.55, 1.32]*	
Moss 2015	AGE	39-41	17.7 ^b	-0.13	0.09	22.2%	0.88 [0.74, 1.04]	
Nystrom 2016	Malmo I	45-70	30.0	-0.13	0.12	13.7%	0.88 [0.70, 1.10]	
Nystrom 2016	Stockholm	40-65	25.0	-0.06	0.17	6.5%	0.94 [0.67, 1.32]	
Subtotal (95%	CI)					71.9%	0.83 [0.75, 0.91]	•
Heterogeneity:	$Tau^2 = 0.00$; $Chi^2 = 3.3$	88, df = 5 (F	$P = 0.64$); $I^2 = 0\%$					
Test for overall	effect: Z = 3.76 (P = 0	.0002)						
Test for subgro	up differences: Chi² =	0.43, df = 1	(P = 0.51); I ² = 09	%).5 0.7 1 1.5 2
aTime since ran	domization; bMedian; '	* Adjusted f	orage: ** Adjuste	d for cluste	erina			1.5 0.7 1 1.5 2 Nammography Usual Care

EVIDENCE SET 1c
Part A- Forest Plot – Breast Cancer Mortality (Short-Case Accrual) (Stratified by Screening Modality)

Reference	Study	Age (at entry)	Mean Follow-up (<u>yrs</u>)	log [RR]	SE	Weight	Risk Ratio [95%CI]	Risk Ratio	
Film Mammog	raphy								
Miller 2014	CNBSS 1 & 2	40-69	21.9	0.05	0.11	15.3%	1.05 [0.85, 1.30]	-	-
Nystrom 2016	Gothenburg	40-59	24.0	-0.30	0.14	9.3%	0.74 [0.56, 0.98]		
Nystrom 2016	Malmo I	45-70	30.0	-0.13	0.12	13.7%	0.88 [0.70, 1.10]		
Nystrom 2016	Malmo II	43-49	22.0	-0.16	0.22	4.0%	0.85 [0.55, 1.32]*		-
Nystrom 2016	Stockholm	40-65	25.0	-0.06	0.17	6.5%	0.94 [0.67, 1.32]		-
Shapiro 1988	HIP	40-64	18.0	-0.24	0.12	12.9%	0.79 [0.63, 1.00]	-	
Subtotal (95%	•					61.6%	0.88 [0.79, 0.98]	•	
Heterogeneity:	$Tau^2 = 0.00$; Chi ²	² = 5.11, df =	5 (P = 0.40); I ² =	2%					
Lest for overall	eπecr / = 2.28 (P = 0.021							
Not Reported	,	,	17 7ª	-0 13	0.09	22 2%	0.88 [0.74, 1.04]	-	
Not Reported Moss 2015	AGE	39-41	17.7ª	-0.13 -0.31	0.09	22.2% 16.2%	0.88 [0.74, 1.04] 0.73 [0.59, 0.90]**	_	
Not Reported Moss 2015 Tabar 2011 St	AGE wedish Two Cou	39-41	17.7³ 29.0⁵	-0.13 -0.31	0.09 0.10	22.2% 16.2% 38.4%	0.88 [0.74, 1.04] 0.73 [0.59, 0.90]** 0.81 [0.67, 0.97]		
Not Reported Moss 2015 Tabar 2011 St Subtotal (95%	AGE wedish Two Cou CI)	39-41 nty 40-74		-0.31		16.2%	0.73 [0.59, 0.90]**	•	
Not Reported Moss 2015 Tabar 2011 S Subtotal (95% Heterogeneity:	AGE wedish Two Cou CI)	39-41 nty 40-74 ² = 1.88, df =	29.0°	-0.31		16.2%	0.73 [0.59, 0.90]**	•	
Subtotal (95% Heterogeneity: Test for overall	AGE wedish Two Cou CI) Tau ² = 0.01; Chi ² effect: Z = 2.28 (39-41 nty 40-74 = 1.88, df = (P = 0.02)	29.0°	-0.31 47%		16.2%	0.73 [0.59, 0.90]**	•	
Not Reported Moss 2015 Tabar 2011 St Subtotal (95% Heterogeneity: Test for overall	AGE wedish Two Cou CI) Tau ² = 0.01; Chi ² effect: Z = 2.28 (39-41 nty 40-74 = 1.88, df = (P = 0.02)	29.0 ^b : 1 (P = 0.17); I ² =	-0.31 47%		16.2%	0.73 [0.59, 0.90]**	0.7 1	1 15 2

EVIDENCE SET 1d
Part A- Forest Plot— Breast Cancer Mortality (Short-Case Accrual) (Stratified by Screening Interval)

Reference	Study	Age (at entry)	Mean Follow-up (<u>yrs)</u>	log [RR]	SE	Weight	Risk Ratio [95%CI]	Risk Ratio (RR) IV, Random, 95% CI
≤ 12 months								
Miller 2014	CNBSS 1 & 2	40-69	21.9	0.05	0.11	15.3%	1.05 [0.85, 1.30]	
Moss 2015	AGE	39-41	17.7ª	-0.13	0.09	22.2%	0.88 [0.74, 1.04]	
Shapiro 1988	HIP	40-64	18.0	-0.24	0.12	12.9%	0.79 [0.63, 1.00]	
Subtotal (95% Heterogeneity:	•	= 3.28. df = 2	(P = 0.19); I ² = 399	%		50.4%	0.90 [0.78, 1.05]	•
	effect: Z = 1.33 (F		. ,					
13-24 months								
Nystrom 2016	Gothenburg	40-59	24.0	-0.30	0.14	9.3%	0.74 [0.56, 0.98]	
Nystrom 2016	Malmol	45-70	30.0	-0.13	0.12	13.7%	0.88 [0.70, 1.10]	
Nystrom 2016	Malmo II	43-49	22.0	-0.16	0.22	4.0%	0.85 [0.55, 1.32]*	•
Subtotal (95% Heterogeneity:		= 0.91, df = 2	(P = 0.63); I ² = 0%			27.0%	0.83 [0.70, 0.97]	•
Test for overall	effect: Z = 2.30 (F	P = 0.02)						
> 24 months								
Nystrom 2016	Stockholm	40-65	25.0	-0.06	0.17	6.5%	0.94 [0.67, 1.32]	
Tabar 2011	Swedish Two County	40-74	29.0 ^b	-0.31	0.10	16.2%	0.73 [0.59, 0.90]**	
Subtotal (95%						22.7%	0.80 [0.63, 1.01]	
Heterogeneity:	Tau ² = 0.01; Chi ²	= 1.56, <u>df</u> = 1	(P = 0.21); I ² = 369	%				
Test for overall	effect: Z = 1.87 (F	P = 0.06)						
Test for subgro	oup differences: C	hi² = 1.01, df =	2 (P = 0.60); I ² = 0)%				
							0.5	0.7 1 1.5 2
		ation; * Adjuste						aphy +/- CBE Usual Care

Appendix 12- Mammography +/- Clincial Breast Exam for Breast-Cancer Mortality (Long-Case Accrual) Forest Plots for Sub-Group Analyses

EVIDENCE SET 2b
Part A- Forest Plot— Breast Cancer Mortality (Long-Case Accrual) (Stratified by use of CBE)

Reference	Study	Age (at entry)	Mean Follow-up (<u>vrs</u>)	log [RR]	SE	Weight	Risk Ratio [95%CI]	Risk Ratio (RR) IV, Random, 95% CI
(+) CBE								
Habbema 1986	HIP	40-64	14.0	-0.25	0.10	16.9%	0.78 [0.64, 0.96]	
Miller 2014	CNBSS 1 & 2	40-69	21.9	-0.01	0.06	21.9%	0.99 [0.88, 1.12]	_
Subtotal (95% C	CI)					38.8%	0.89 [0.71, 1.12]	
Heterogeneity: 7	Tau ² = 0.02; Chi ²	= 3.85, df =	$1 (P = 0.05); I^2 = 7$	74%				
Test for overall e	effect: Z = 0.97 (P = 0.33)						
(-) CBE								
Bjurstam 2003	Gothenburg	40-59	13.8	-0.29	0.13	14.0%	0.75 [0.58, 0.97]	
Moss 2015	AGE	39-41	17.7a	-0.07	0.08	19.9%	0.93 [0.80, 1.09]	-
Tabar 1995	Kopparberg	40-74	12.5	-0.51	0.14	13.4%	0.60 [0.46, 0.79]*	
Tabar 1995	Ostergotland	40-74	12.5	-0.25	0.13	13.9%	0.78 [0.60, 1.01]*	
Subtotal (95% (CI)					61.2%	0.77 [0.64, 0.93]	
Heterogeneity: 7	Tau ² = 0.02; Chi ²	= 8.22, df =	$3 (P = 0.04); I^2 = 6$	64%				
Test for overall e	effect: Z = 2.67 (P = 0.008)						
Test for subgrou	p differences: C	hi² = 0.91, d	f = 1 (P = 0.34); I ²	= 0%				
· ·	•	,	. ,,					
								0.5 0.7 1 1.5 2
Median: * Adius	sted for age and	clustering					1	Mammography Usual Care

EVIDENCE SET 2c
Part A- Forest Plot – Breast Cancer Mortality (Long-Case Accrual) (Stratified by Screening Modality)

Reference	Study	Age (at entry)	Mean Follow-up (<u>vrs</u>)	log [RR]	SE	Weight	Risk Ratio [95%CI]	Risk Ratio (RR) IV, Random, 95% CI
Film Mammogr	aphy							
Bjurstam 2003	Gothenburg	40-59	13.8	-0.29	0.13	14.0%	0.75 [0.58, 0.97]	
Habbema 1986	HIP	40-64	14.0	-0.25	0.10	16.9%	0.78 [0.64, 0.96]	
Miller 2014	CNBSS 1 & 2	40-69	21.9	-0.01	0.06	21.9%	0.99 [0.88, 1.12]	-
Subtotal (95%	CI)					52.8%	0.85 [0.70, 1.03]	
Heterogeneity:	Tau ² = 0.02; Chi	² = 6.19, df	= 2 (P = 0.05); I ² =	- 68%				
Test for overall	effect: Z = 1.62	(P = 0.11)						
Not Reported Moss 2015	AGE	39-41	17.7ª	-0.07	0.08	19.9%	0.93 [0.80, 1.09]	
Tabar 1995	Kopparberg	40-74	12.5	-0.51	0.14	13.4%	0.60 [0.46, 0.79]* —	
Tabar 1995	Ostergotland	40-74	12.5	-0.25	0.13	13.9%	0.78 [0.60, 1.01]*	<u> </u>
Subtotal (95%	•••••	40-74	12.0	-0.23	0.10	47.2%	0.77 [0.60, 1.00]	
•	•	² = 7.81. df	= 2 (P = 0.02); I ² =	= 74%				
Test for overall		•	, ,,					
		,						
Test for subgrou	up differences: ($Chi^2 = 0.37,$	df = 1 (P = 0.54);	2 = 0%				
aMedian; *Adjus	sted for age and	clustering					0. Mammo	5 0.7 1 1.5 2 graphy +/- CBE Usual Care

EVIDENCE SET 2d
Part A- Forest Plot – Breast Cancer Mortality (Long-Case Accrual) (Stratified by Screening Interval)

Reference	Study	Age (at entry)	Mean Follow-up (<u>vrs</u>)	log [RR]	SE	Weight	Risk Ratio [95%CI]	Risk Ratio (RR) IV, Random, 95% CI
≤12 months							0.70 (0.04, 0.00)	
Habbema 1986	HIP	40-64	14.0	-0.25	0.10	16.9%	0.78 [0.64, 0.96]	
Miller 2014	CNBSS 1 & 2	40-69	21.9	-0.01	0.06	21.9%	0.99 [0.88, 1.12]	-
Moss 2015	AGE	39-41	17.7ª	-0.07	0.08	19.9%	0.93 [0.80, 1.09]	
Subtotal (95% C	I)					58.7%	0.92 [0.81, 1.04]	•
Heterogeneity: Ta	au ² = 0.01; Chi ²	e = 3.85, df =	2 (P = 0.15); l ² = 4	8%				
Test for overall ef	ffect: Z = 1.39 (P = 0.16)						
13-24 months Bjurstam 2003	Gothenburg	40-59	13.8	-0.29	0.13	14.0%	0.75 [0.58, 0.97]	
Subtotal (95% C	_	40-33	10.0	-0.23	0.10	14.0%	0.75 [0.58, 0.97]	
Heterogeneity: N	ot applicable							
Test for overall e	ffect: Z = 2.19	(P = 0.03)						
>24 months								
Tabar 1995	Kopparberg	40-74	12.5	-0.51	0.14	13.4%	0.60 [0.46, 0.79]*	-
Tabar 1995 C	Ostergotland	40-74	12.5	-0.25	0.13	13.9% 27.3%	0.78 [0.60, 1.01]* 0.69 [0.53, 0.89]	
Heterogeneity: Ta	au ² = 0.02; Chi ²	2 = 1.88, df =	= 1 (P = 0.17); I ² = 4	17%				
Test for overall ef	ffect: Z = 2.87 (P = 0.004)						
T		NE:2 404	# 0 (D 0 00) I	FO 00/				
rest for subgroup	anterences: C	/III~ = 4.91, ($df = 2 (P = 0.09); l^2$	= 59.3%				
								0.5 0.7 1 1.5 2
aMedian; *Adjuste	ed for age and	clustering					Mam	nmography +/- CBE Usual Care

Appendix 13- Mammography +/- Clincial Breast Exam for All-Cause Mortality - Forest Plots for Sub-Group Analyses

EVIDENCE SET 3b

Part A- Forest Plot – All-Cause Mortality (Stratified by CBE)

Reference	Study	Age	Mean	log	SE	Weight	Risk Ratio	Risk Ratio (RR) IV, Random, 95% CI	
(+) CBE	Otaay	(at entry)	Follow-up (vrs)	[RR]		Weight	[95%CI]	IV, Random, 95% CI	
Aron and Prorok 1	986 HIP	40-64	10.0	-0.01	0.03	4.7%	0.99 [0.93, 1.05]	_	
Miller 2014	CNBSS 1 & 2			0.02	0.03	11.4%	1.02 [0.98, 1.06]	1	
Subtotal (95% CI)		40-59	25.0	0.02	0.02	16.1%	1.01 [0.98, 1.04]		
		100 df = 4 /	D = 0.40\: 12 = 00/			10.176	1.01 [0.00, 1.04]	Y	
Heterogeneity: Tau	•	,	P = 0.42); P = 0%						
Test for overall effe	ect: Z = 0.65 (P =	0.51)							
(-) CBE									
Moss 2015	AGE	39-41	17.7a	-0.02	0.03	6.7%	0.98 [0.93, 1.03]	+	
Nystrom 2002	Gothenburg	40-59	13.2	-0.06	0.03	4.3%	0.94 [0.88, 1.00]	-	
Nystrom 2002	Ostergotland	40-74	17.2	-0.01	0.01	42.8%	0.98 [0.95, 1.01]	•	
Nystrom 2002	Stockholm	40-64	14.7	0.03	0.08	0.8%	0.99 [0.95, 1.03]	+	
Nystrom 2002	Malmo I	45-70	19.2	-0.02	0.02	18.6%	0.99 [0.97, 1.01]		
Nystrom 2002	Malmo II	43-49	9.1	-0.01	0.02	10.7%	1.03 [0.89, 1.20]		
Subtotal (95% CI))					83.9%	0.98 [0.97, 1.00]	•	
Heterogeneity: Tau	u² = 0.00; Chi² = 2	.84, df = 5 (P = 0.72); I ² = 0%					Ì	
Test for overall effe	ect: Z = 2.09 (P =	0.04)							
Test for subgroup	differences: Chi ²	= 2.07, df =	1 (P = 0.15); I ² = 51.	.6%					
							0.5	0.7 1 1.5	_
^a Median							Mami	mography Usu	ual Care

EVIDENCE SET 3c
Part A- Forest Plot – All-Cause Mortality (Stratified by Screening Modality)

Reference	Study	Age (at entry) F	Mean Follow-up (yrs)	log [RR]	SE	Weight	Risk Ratio [95%CI]	Risk Ratio (RR) IV, Random, 95% CI	
Film Mammography									
Aron and Prorok 1986	HIP	40-59	10.0	-0.01	0.03	4.7%	0.99 [0.93, 1.05]	+	
Miller 2014	CNBSS 1 & 2	40-59	25.0	0.02	0.02	11.4%	1.02 [0.98, 1.06]	+	
Nystrom 2002	Gothenburg	40-59	13.2	-0.06	0.03	4.3%	0.94 [0.88, 1.00]		
Nystrom 2002	Malmo I	45-70	19.2	-0.01	0.01	42.8%	0.99 [0.97, 1.01]	•	
Nystrom 2002	Malmo II	43-49	9.1	0.03	0.08	0.8%	1.03 [0.89, 1.20]		
Nystrom 2002 Subtotal (95% CI)	Stockholm	40-64	14.7	-0.01	0.02	10.7% 74.7%	0.99 [0.95, 1.03] 0.99 [0.98, 1.01]	ţ	
Heterogeneity: Tau ² =	0.00; Chi ² = 4.	95, df = 5 (P	= 0.42); I ² = 0%						
Test for overall effect:	Z = 1.03 (P =	0.30)							
Not Reported Moss 2015	AGE	39-41	17.7ª	-0.02	0.03	6.7%	0.98 [0.93, 1.03]	+	
Nystrom 2002 Subtotal (95% CI)	Ostergotland	40-74	17.2	-0.02	0.02	18.6% 25.3%	0.98 [0.95, 1.01] 0.98 [0.95, 1.01]	•	
Heterogeneity: Tau ² =	0.00; Chi ² = 0.	00, df = 1 (P	= 1.00); I ² = 0%						
Test for overall effect:	Z = 1.51 (P = 0	0.13)	,						
Test for subgroup diffe	rences: Chi² =	0.61, df = 1	(P = 0.43); I ² = 0%						
								0.5 0.7 1 1.5 2	
ªMedian								ography +/- CBE Usual Care	

EVIDENCE SET 3d Part A- Forest Plot – All-Cause Mortality (Stratified by Screening Interval)

Reference	Study	Age (at entry)	Mean Follow-up (<u>yrs</u>)	log [RR]	SE	Weight	Risk Ratio [95%CI]	Risk Ratio (RR) IV, Random, 95% CI
≤12 months								
Aron and Prorok 1986	HIP	40-59	10.0	-0.01	0.03	4.7%	0.99 [0.93, 1.05]	+
Miller 2014 C	NBSS 1 & 2	40-59	25.0	0.02	0.02	11.4%	1.02 [0.98, 1.06]	+
Moss 2015	AGE	39-41	17.7ª	-0.02	0.03	6.7%	0.98 [0.93, 1.03]	+
Subtotal (95% CI)						22.8%	1.00 [0.97, 1.03]	♦
Heterogeneity: Tau ² =	0.00; Chi ² =	1.67, df = 2	(P = 0.43); $I^2 = 0\%$					
Test for overall effect:	Z = 0.13 (P =	= 0.90)						
13-24 months								
Nystrom 2002	Gothenburg	40-59	13.2	-0.06	0.03	4.3%	0.94 [0.88, 1.00]	-
Nystrom 2002	Malmo I	45-70	19.2	-0.01	0.01	42.8%	0.99 [0.97, 1.01]	•
Nystrom 2002	Malmo II	43-49	9.1	0.03	0.08	0.8%	1.03 [0.89, 1.20]	-
Subtotal (95% CI)						47.9%	0.98 [0.95, 1.01]	♦
Heterogeneity: Tau ² =	0.00; Chi ² =	2.63, df = 2	(P = 0.27); I ² = 24%	5				
Test for overall effect:	Z = 1.12 (P =	= 0.26)						
>24 months								
Nystrom 2002 Q	stergotland	40-74	17.2	-0.02	0.02	18.6%	0.98 [0.95, 1.01]	•
Nystrom 2002	Stockholm	40-64	14.7	-0.01	0.02	10.7%	0.99 [0.95, 1.03]	+
Subtotal (95% CI)						29.3%	0.98 [0.96, 1.01]	♦
Heterogeneity: Tau ² =	0.00; Chi ² =	0.15, df = 1	(P = 0.69); I ² = 0%					
Test for overall effect:	Z = 1.32 (P =	= 0.19)						
Test for subgroup diffe	erences: Chi²	= 1.26, df =	= 2 (P = 0.53); I ² = 0°	%				
							0.5	0.7 1 1.5 2
^a Median							V.5 Mammograp	

Appendix 14- Extracted False Positive Studies (Overview of Reviews)

False Positive Recalls and False Positive Biopsies – Mammography vs Usual Care

Study (Review)	Country; Source of data	Details	False Positive Recalls	False Positive Biopsies
Hubbard (40) (USPSTF 2016)	US; Breast Cancer Surveillance Consortium (BSCS)	Data from 7 BCSC centres from 1994-2006	Cumulative FP recall rate after 10 years (%, 95%CI) for annual and biennial screening, respectively: • Ages 40-49: 61.3% (59.4-63.1%) and 41.6% (40.6-42.5%) • Ages 50-59: 61.3% (58.0-64.75%) and 42.05 (40.4-43.7%)	Cumulative probability of FP biopsies after 10 years (% and 95% CI), for annual and biennial rates, respectively: • Ages 40-49: 7.0% (6.1-7.8%) and 4.8% (4.4-5.2%) • Ages 50-59, 9.4% (7.4-11.5%) and 6.4% (5.6-7.2)
Kerlikowske et al., 2013 (41) (USPSTF 2016 [recall]; USPSTF 2016 and ACS 2014 [biopsies])	US; BCSC	Data 7 BCSC centres from 1994-2008	Cumulative probability of FP mammogram, women aged 40-74, first stratified by age, then by frequency of screening, then by breast density category: Ages 40-49 • Generally, highest for annual screening interval, followed by biennial, with triennial having the lowest FP rate • Fatty and scattered breast density had lower FP rates compared to heterogeneous and extreme Ages 50-74 • Above patterns are also observed Women aged 40-49 generally had higher rates of FP compared to 50-74.	 FP biopsy rate data stratified by age, then by screening interval, and then by breast density: Similar overall patterns as was seen in FP recall For ages 40-49, the screening interval with the highest FP biopsy rate was annual, followed by biennial, then triennial Heterogeneous and extreme breast density had higher rates of FP biopsies compared to fatty and scattered. Higher FP biopsy rates were observed for 40-49 compared to 50-74
Unpublished data (USPSTF 2016)	US; BCSC	6 BCSC centres from 2003-2011	FP rate per 1,000 women screened (95%CI)	Unclear whether the data provided for biopsies were specific to patients who had FP results

Study (Review)	Country; Source of data	Details	False Positive Recalls	False Positive Biopsies
			higher rates of FP for the 9-18 mo In contrast, ages 60-69 and 70-79 had higher rates of FP in the 19-30 months. The p-values for all comparisons were not statistically significant(1).	
			By screening interval, 11-14 mo vs 23-26 mo: • All age categories had higher rates of FP for 11-14 mo except for 60-69. p-values for all comparisons not statistically significant	
			all age categories generally had higher FP rates for heterogeneous and extreme breast density compared to fatty-scattered, Except 70-79, where extreme breast density had a lower FP rate compared to fatty-scattered. p-values for all comparisons were statistically significant.	
			For race: • General pattern: for all categories higher FP rate for Whites, followed by Hispanics, Blacks, and then Asians. • The 'Other' category generally had a high FP rate comparable to the 'Whites' category. All comparisons were statistically significant except 60-69	
Elmore et al., 1998 (42) (USPSTF 2016, CTFPHC 2011 [recall]; USPSTF 2016 [biopsies])	US; Not specified	Data from 11 breast cancer screening centres from 1983-1995	Overall cumulative risk of a FP (% and 95%CI) after 10 screening mammograms: • 49% (40.3-64.1%). • Ages 40-49: 56% (39.5-75.8%) • Ages 50-59: 47% (37.8-63.0%)	Overall rate was 19% (9.8-41.2%) for at least one FP biopsy.
Hofvind et al., 2004 (43) (CTFPHC 2011	Norway; Norwegian Breast Cancer Screening	No additional information reported	For women aged 50-51 who participated in 3 biennial screening rounds, the FP recall rate during period of 20 years was 20.8%	20 year cumulative FP biopsy rate (% and 95% CI) • Ages 50-59: 4.1% (3.9-4.3%)

Study (Review)	Country; Source of data	Details	False Positive Recalls	False Positive Biopsies
as reported in USPSTF 2009; not reported in USPSTF 2016) – recall Roman et al., 2013 (47) (ACS	Program			
Malmo (ACS 2014)	Sweden; Mammography RCT		FP rate of 1.26% in the mammography group	
Stockholm (ACS 2014)	Sweden; Mammography RCT		355 FPs out of 100,000 woman-years for the mammography group	NR
Schonberg et al., 2009 (46) (ACS 2014)	US; Not specified	Cohort study	Women 80 years and older FP rate in the screened group was 10.64%	FP biopsy rate of 1.84%

False Positive Recalls and False Positive Biopsies - Clinical Breast Exam vs Usual Care

Study (Review)	Country; Source of data	Details	False Positive Recalls	False Positive Biopsies
Abuidris et al., 2013 (56) (ACS 2014)	Sudan; RCT	No additional information	FP rate of 0.9% for receiving CBE once compared to no screening.	NR
Sankarana-Rayanan et al., 2011 (57) (ACS 2014)	India; RCT	No additional information	FP rate of 5.7% for receiving CBE (every 3 years) compared to no screening (5.5-5.9%)	NR
Pisani et al., 2006 (CTPFHC 2011 as reported in USPSTF 2009 [but not USPSTF 2016])	Unknown; RCT	No additional information	No results reported in systematic review	NR

False Positive Biopsies (Unnecessary Biopsies) - Breast Self Exam vs No Screening

Study (Review)	Country;	; Details False Positive Reca		False Positive Biopsies
	Source of data			
Semiglazov et al., 2003	Russia;	No additional information	NR	Benign biopsy rate of RR 2.05 (1.80-2.33)
(CTFPHC 2011)	Cluster RCT			
Thomas et al., 2002 (CTFPHC	China;	No additional information	NR	Benign biopsy rate of RR 1.57 (1.48-1.68)
2011)	Cluster RCT			

Appendix 15. List of potentially relevant, unpublished RCTs

Trial Identifier	Study Title	Estimated Study Completion Date
	Initial Evaluation of Ultra FAST Breast Magnetic Resonance in Breast	
NOT0000 400 4	Cancer Screening: Comparative Study With Mammography and	February 2017 (not
NCT02324894	Ultrasound.	available)
NCT02306265	Assessment of Diagnostic Accuracy and Performance of Digital Breast Tomosynthesis Compared to Mammography (ADAPT)	July 2017
NCT02777164	Evaluation of a Three Dimensional Functional Metabolic Imaging and Risk Assessment System for Classifying Women at High Risk of Breast Cancer	August 2017
NCT02155075	Evaluation of REAL IMAGING'S 3D Functional Metabolic Imaging and Risk Assessment ("3D MIRA") System in Women at High Risk for Breast Cancer	August 2017
	The Assessment of the Role of Automated Breast Ultrasound (ABUS) in Screening Women With Dense Breasts for Early Detection of Breast	- C
NCT02386176	Cancer	November 2017
NCT01091545	Malmö Breast Tomosynthesis Screening Trial (MBTST)	December 2017
NCT02066142	Tomosynthesis (TS) Versus Ultrasonography (US) in Women With Dense Breast (ASTOUND)	July 2018
NCT02698202	Screening for Breast Cancer With Digital Breast Tomosynthesis	December 2018
NCT02033486	Digital Breast Tomosynthesis Guided Tomographic Optical Breast Imaging (TOBI)	January 2019
NCT02616432	Tomosynthesis Mammography Imaging Screening Trial (TMISTLead-in)	November 2019
NCT02933489	Abbreviated Breast MRI and Digital Tomosynthesis Mammography in Screening Women With Dense Breasts	December 2019
NCT01315015	Breast Cancer Screening With MRI in Women Aged 50-75 Years With Extremely Dense Breast Tissue: the DENSE Trial	December 2019
NCT02590315	Tomosynthesis Versus Digital Mammography in a Population-based Screening Program (ProteusDonna)	December 2019
NCT02835625	The Tomosynthesis Trial in Bergen (TOBE)	January 2022
NCT02643966	Assessment of Periodic Screening of Women With Denser Breast Using WBUS and DBT (DBTUST)	December 2022
ISRCTN33292440	Nationwide cluster-randomised trial of extending the NHS breast screening age range in England	December 2026
NCT02210546	Contrast-enhanced MR Imaging as a Breast Cancer Screening in Women at Intermediate Risk (MRIB)	Unknown

NCT00971087	Multicenter Hologic Tomosynthesis Study	Unknown
	Efficacy of contrast enhanced spectral mammography versus standard of	
	care imaging tests (tomosynthesis and ultrasound) in women with	
	mammographically dense breast tissue recalled for investigation of	
ACTRN12616000533493	abnormalities detected on routine screening mammograms	Unknown
	Early detection of breast cancer by self examination, clinical examination	
	and fine needle aspiration cytology in rural women -a population based	
CTRI/2016/04/006865	study	Unknown

Appendix 16: Evaluation of Subgroup analyses (GRADE Criteria)

Based on GRADE criteria (BMJ 2010; JAMA 2014)

Subgroup variables:

• Age, ethnicity, SES, geographic location, breast density, screening interval, advancements in screening technology (film, digital, etc), type of control (no screening vs usual care)

Additional guiding points: should be skeptical when evidence at very high risk of bias; subgroup effects exist along a continuum, not a 'accept or reject' situation.

Criteria	Explanation	Assessment
1. Is the subgroup	More credible when variables defined	Yes.
variable a	at time of randomization. The	All based on assessments at
characteristic	credibility of subgroup hypotheses	baseline (or prespecified,
specified at	based on post-randomization	such as screening interval).
baseline?	characteristics is severely	
	compromised, and can be rejected	
	simply on this criterion.	
2. Is the subgroup	Between-study comparisons are	No (all except age- The AGE
difference suggested	limited because a number of	trial only contributed 39-41
by comparisons	competing explanations can explain	age group data 'between',
within rather than	the results. Within-trial subgroup	whereas other studies
between studies.	differences are stronger. Most	provided data for multiple
	subgroup analyses from systematic	age groups- 'within').
	reviews are limited by between-study	
	comparisons.	'Yes' answer based on a mix
		of between and within study
		comparisons and results are
		consistent across studies

3. Does statistical analysis suggest that chance is an unlikely explanation for the subgroup difference?	Need to look at degree of overlap of confidence intervals between subgroups. Would also apply if confidence intervals are substantially overlapping when point estimates differ. Check test of interaction.	No. Substantial overlap of subgroups. Test for subgroup differences are not statistically significant.
4. Did the hypothesis precede rather than follow the analysis and include a hypothesized direction that was subsequently confirmed?	Credibility of post hoc hypotheses is questionable. Multiple comparisons issue. Specification of direction of effect a priori. Failure to correctly identify the direction of subgroup effect will weaken the inference.	Yes, but direction was not prespecified.
5. Was the subgroup hypothesis one of a small number tested?	Strength of inference for confirmation of any hypothesis will decrease in a large number of hypotheses are tested.	No, a moderate number of subgroup hypotheses were pre-specified.
6. Is the subgroup difference consistent across studies?	Replication in other studies increases credibility.	No subgroup difference; consistent results across studies.
7. Does external evidence (biological or sociological rationale) support the hypothesized	Does additional, external evidence exist to support the subgroup claim? Would need to be strong. Are the subgroup differences	Is there other, relevant evidence that would lead one to believe that there might be subgroup differences for age?
subgroup difference?	challenged by current biological (or other) understanding?	All others – no evidence exists (unknown)

Appendix 17: False Positive Calculations

	40-49											
	CTFPHC 2011 (Using 2005- 2006) [Initial+ 3(Subsequ ent)]	A (SC) (2011- 2012) Used method from CTFPHC 2011 [initial + 7(subseq uent)]	B (LC) (2011- 2012) Used method from CTFPHC 2011 [initial + 4(subseq uent)]	C (2005- 2006) Treated data as cross- sectional . Initial screen data	D (2005- 2006) Treated data as cross- sectional. Subseque nt screen data	E (SC) (2011- 2012) Treated data as cross- sectional. Initial screen data	F (SC) (2011-2012) Treated data as cross- sectional. Subsequent screen data.	G (LC) (2011-2012) Treated data as cross- sectional. Initial screen data	H (LC) Treated data as cross- sectional. Subsequent screen data	I (2005-2006) Treated data as cross- sectional. Initial + Subsequent (weighted average)	J (SC) (2011-2012) Treated data as cross- sectional. Initial + Subsequen t (weighted average)	K (LC) (2011-2012) Treated data as cross- sectional. Initial + Subsequen t (weighted average)
	omen screene											
FP Mam.	327	660	442	134	64	148	73	148	73	86	92	92
Un. biopsies	36	90	64	19	6	28	9	28	9	10	14	14
Per one brea	ast cancer dea	th prevented	d						<u> </u>			
NNS	2,108	2,000	3,704									
FP Mam.	690	1,320	1,639									
Un. biopsies	75	180	242									

	50-59											
	CTFPHC 2011 (Using 2005- 2006) [Initial+ 3(Subsequ ent)]	A (SC) (2011- 2012) Used method from CTFPHC 2011 [initial + 7(subseq uent)]	B (LC) (2011- 2012) Used method from CTFPHC 2011 [initial + 4(subseq uent)]	C (2005-2006) Treated data as cross-sectional . Initial screen data	D (2005- 2006) Treated data as cross- sectional. Subsequ ent screen data	E (SC) (2011-2012) Treated data as cross- sectional. Initial screen data	F (SC) (2011-2012) Treated data as cross- sectional. Subsequent screen data	G (LC) (2011- 2012) Treated data as cross- sectional. Initial screen data	H (LC) Treated data as cross- sectional. Subseque nt screen data	I (2005- 2006) Treated data as cross- sectional. Initial + Subsequ ent (weighte d average)	I (SC) (2011- 2012) Treated data as cross- sectional. Initial + Subsequ ent (weighte d average)	J (LC) (2011- 2012) Treated data as cross- sectional. Initial + Subsequ ent (weighte d average)
Per 1,000 w	omen screene	d										
FP Mam.	NR	652	437	122	58	151	73	151	73	77	90	90
Un. biopsies	NR	80	55	17	7	21	9	21	9	10	12	12
Per one brea	ast cancer dea	th prevented	d									
NNS	NR	1,136	962									
FP Mam.	NR	741	420									
Un. biopsies	NR	91	53									

60-69												
Par 1 000 w	CTFPHC 2011 (Using 2005- 2006) [Initial+ 3(Subsequ ent)]	A (SC) (2011- 2012) Used method from CTFPHC 2011 [initial + 7(subseq uent)]	B (LC) (2011- 2012) Used method from CTFPHC 2011 [initial + 4(subseq uent)]	C (2005- 2006) Treated data as cross- sectional . Initial screen data	D (2005- 2006) Treated data as cross- sectional. Subsequ ent screen data	E (SC) (2011-2012) Treated data as cross- sectional. Initial screen data	F (SC) (2011- 2012) Treated data as cross- sectional. Subseque nt screen data	G (LC) (2011- 2012) Treated data as cross- sectional. Initial screen data	H (LC) Treated data as cross- sectional. Subseque nt screen data	I (2005-2006) Treated data as cross- sectional. Initial + Subsequent (weighted average)	J (SC) (2011-2012) Treated data as cross- sectional. Initial + Subsequent (weighted average)	K (LC) (2011-2012) Treated data as cross- sectional. Initial + Subsequent (weighted average)
	Per 1,000 women screened											
FP Mam.	NR	578	385	95	51	128	64	128	64	56	69	69
Un. biopsies	NR	76	51	13	7	18	8	18	8	8	9	9
Per one brea	Per one breast cancer death prevented											
NNS	NR	541	452									
FP Mam.	NR	312	174									
Un. biopsies	NR	41	23									

70-74												
	CTFPHC 2011 (Using 2005- 2006)	A (SC) (2011- 2012) Used method from CTFPHC 2011 [initial + 3(subseq uent)]	B (LC) (2011- 2012) Used method from CTFPHC 2011 [initial + 3(subseq uent)]	C (2005- 2006) Treated data as cross- sectional . Initial screen data	D (2005- 2006) Treated data as cross- sectional. Subseque nt screen data	E (SC) (2011- 2012) Treated data as cross- sectional. Initial screen data	F (SC) (2011- 2012) Treated data as cross- sectional. Subseque nt screen data	G (LC) (2011-2012) Treated data as cross- sectional. Initial screen data	H (LC) Treated data as cross- sectional. Subsequent screen data	I (2005-2006) Treated data as cross- sectional. Initial + Subsequent (weighted average)	J (SC) (2011-2012) Treated data as cross- sectional. Initial + Subsequent (weighted average)	K (LC) (2011-2012) Treated data as cross-sectional. Initial + Subsequent (weighted average)
Per 1,000 women screened												
FP Mam.	212	274	274	80	44	109	55	109	55	47	58	58
Un. biopsies	26	68	38	10	5	15	8	15	8	6	7	7
Per one brea	Per one breast cancer death prevented											
NNS	451	885	699									
FP Mam.	96	438	230									
Un. biopsies	11	60	31									

Appendix 18: Organized Breast Cancer Screening Programs

Commencement of Organized Screening Programs								
	Canada	UK	USA	Sweden				
Trial	CNBSS 1&2	AGE	HIP	Malmo I, Malmo II, Stockholm, Gothenburg, Swedish Two Counties				
Start Year	1980	1991	1963	1976-1982				
Age at Entry	40-59	39-41	40-64	39-74				
Screening Duration	5 years	8 years	3 years	4-12 years				
Longest Follow-up	21.9 yrs (mean)	17.7 yrs (median)	18 yrs (mean)	22-30 yrs (mean)				
Start Year (organized screening)	1988	1988	1991	1986				
Age	50-69	50-70	NR	40-74				
Technology surveyed in 2007-2008	Film, digital, CBE	Film, digital	NR	NR				
% of population covered in 1995	(50-69): <25% In 2014 (50-69): 54.1% In 2013 (50-69): 53.9% In 2010 (50-69): 53.2% In 2009 (50-69): 52.1% *Reported as 47.3% in previous iteration of report. In 2008 (50-69): 45.9%	100%	25-50%	100%				

NR: not reported.

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