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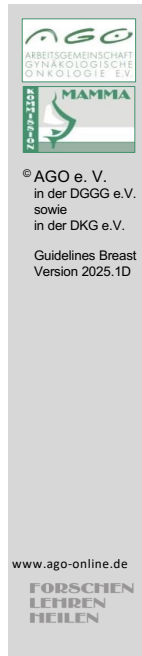
Guidelines Breast  
Version 2025.1D

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# Diagnostik und Therapie früher und fortgeschrittener Mammakarzinome

## Früherkennung und Diagnostik

# Früherkennung und Diagnostik



- **Versionen 2005 - 2024:**

**Albert / Blohmer / Fallenberg / Fersis / Gerber / Heil / Junkermann /  
Kühn / Maass / Müller-Schimpfle / Scharl / Schreer / Wöckel**

- **Version 2025:**

**Fallenberg / Thomssen**

## Screened data bases

Pubmed        2018 - 2024  
Medline       2018 - 2024  
Cochrane      2018 - 2024

## Guidelines

S3 Diagnostik, Therapie und Nachsorge des Mammakarzinoms:

1. Wöckel A, Festl J, Stüber T et al. Interdisciplinary Screening, Diagnosis, Therapy and Follow-up of Breast Cancer. Guideline of the DGGG and the DKG (S3-Level, AWMF Registry Number 032/045OL, December 2017) - Part 1 with Recommendations for the Screening, Diagnosis and Therapy of Breast Cancer. Geburtshilfe Frauenheilkd. 2018 Oct;78(10):927-948. doi: 10.1055/a-0646-4522. Epub 2018 Oct 19.
2. Wöckel A, Festl J, Stüber T et al. Interdisciplinary Screening, Diagnosis, Therapy and Follow-up of Breast Cancer. Guideline of the DGGG and the DKG (S3-Level, AWMF Registry Number 032/045OL, December 2017) - Part 2 with Recommendations for the Therapy of Primary, Recurrent and Advanced Breast Cancer. Geburtshilfe Frauenheilkd. 2018 Nov;78(11):1056-1088. doi: 10.1055/a-0646-4630. Epub 2018 Nov 26.

European Commission Initiative on Breast Cancer (ECIBC)

European guidelines on breast cancer screening and diagnosis

<https://healthcare-quality.jrc.ec.europa.eu/european-breast-cancer-guidelines>


2015 ACS Update Breast Cancer Screening for women at average risk

IARC Handbook 2016

European Commission 2016

(<http://ecibc.jrc.ec.europa.eu/recommendations/list/3>; Update 24.11.2016, Abruf 20122016)

Screened: Metaanalyses/ Systematic reviews / RCT / Cohort studies



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## Früherkennung bei asymptomatischen Frauen durch Mammographie

<u>Alter</u>	<u>Intervall (Monate)</u>	<u>Oxford</u>		
		<u>LOE</u>	<u>GR</u>	<u>AGO</u>
• < 40	na	-	-	--
• 40-44				
• normales Risiko	na	1b	B	-
• moderat erhöhtes Risiko oder Eigenanamnese*	na	1b	B	+/-
• 45-49 <sup>#</sup>	24-36	1a	A	+ <sup>#</sup>
• 50-75 <sup>**</sup>	24	1a	A	++
• > 75 <sup>***</sup>	24	4	C	+/-

\* Siehe Empfehlungen der Strahlenschutzkommission, außer INFP (intensiviertes Früherkennungsprogramm) siehe Kapitel 02.

\*\* Nationales Mammographie-Screening-Programm.

\*\*\* Abhängig von Gesundheitszustand + Lebenserwartung (> 10 Jahre).

# Rechtfertigende Indikation ist notwendig, solange Bundesmantelvertrag (Screening) nicht angepasst.

1. Bennett A, Shaver N, Vyas N et al. Screening for breast cancer: a systematic review update to inform the Canadian Task Force on Preventive Health Care guideline. Syst Rev. 2024 Dec 19;13(1):304.
2. European Commission Initiative on Breast Cancer (ECIBC): European guidelines on breast cancer screening and diagnosis (<https://cancer-screening-and-care.jrc.ec.europa.eu/en/ecibc/european-breast-cancer-guidelines>) Download 15.01.2025
3. Spear G, Lee K, DePersia A et al. Updates in Breast Cancer Screening and Diagnosis. Curr Treat Options Oncol. 2024 Nov;25(11):1451-1460.
4. Thomassin-Naggara I, Kilburn-Toppin F, Athanasiou A et al; EUSOBI Board. Misdiagnosis in breast imaging: a statement paper from European Society Breast Imaging (EUSOBI)-Part 1: The role of common errors in radiology in missed breast cancer and implications of misdiagnosis. Eur Radiol. 2024 Nov 15.
5. Thomassin-Naggara I, Kilburn-Toppin F, Athanasiou A et al; EUSOBI Board. Misdiagnosis in breast imaging: a statement paper from European Society Breast Imaging (EUSOBI)-Part 2: Main causes of errors in breast imaging and recommendations from European Society of Breast Imaging to limit misdiagnosis. Eur Radiol. 2024 Nov 15.
6. Trentham-Dietz A, Chapman CH, Jayasekera J et al. Breast Cancer Screening With Mammography: An Updated Decision Analysis for the U.S. Preventive Services Task Force [Internet]. Rockville (MD): Agency for Healthcare Research and Quality (US); 2024 Apr.

Report No.: 23-05303-EF-2.

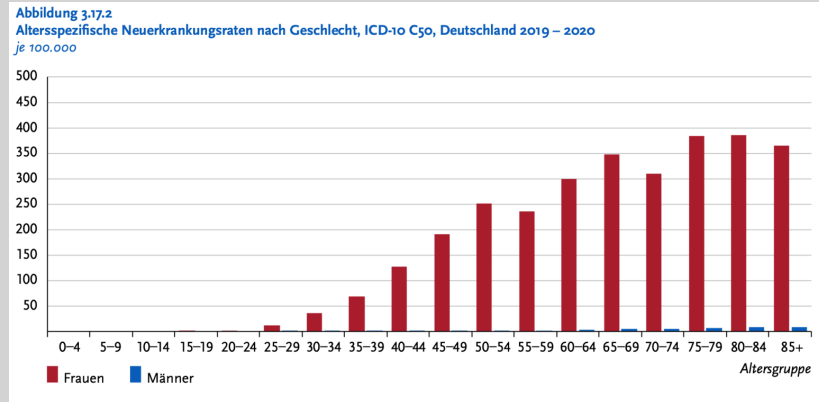
7. Jayasekera J, Stein S, Wilson OWA et al. Benefits and Harms of Mammography Screening in 75 + Women to Inform Shared Decision-making: a Simulation Modeling Study. *J Gen Intern Med.* 2024 Feb;39(3):428-439.
8. Expert Panel on Breast Imaging; Brown A, Lourenco AP, Niell BL et al. ACR Appropriateness Criteria® Transgender Breast Cancer Screening. *J Am Coll Radiol.* 2021 Nov;18(11S):S502-S515.
9. Monticciolo DL, Malak SF, Friedewald SM et al. Breast Cancer Screening Recommendations Inclusive of All Women at Average Risk: Update from the ACR and Society of Breast Imaging. *J Am Coll Radiol.* 2021 Sep;18(9):1280-1288.
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11. Schünemann HJ, Lerda D, Quinn C et al. Breast Cancer Screening and Diagnosis: A Synopsis of the European Breast Guidelines. *Annals of Internal Medicine.* 2020;172(1):46-56.
12. Zielonke N, Kregting LM, Heijnsdijk EAM et al. The potential of breast cancer screening in Europe. *International journal of cancer Journal international du cancer.* 2021;148(2):406-18.
13. Maroni R, Massat NJ, Parmar D et al. Screening Guidelines Update for Average-Risk and High-Risk Women. *AJR American journal of roentgenology.* 2020;214(2):316-23.
14. Mao Z, Nystrom L, Jonsson H. Breast cancer screening with mammography in women aged 40-49 years: Impact of length of screening interval on effectiveness of the program. *Journal of medical screening.* 2020:969141320918283.
15. Khil L, Heidrich J, Wellmann I et al. Incidence of advanced-stage breast cancer in regular participants of a mammography screening program: a prospective register-based study. *Bmc Cancer.* 2020;20(1):174.
16. Duffy SW, Tabar L, Yen AM, et al. Mammography screening reduces rates of advanced and fatal breast cancers: Results in 549,091 women. *Cancer.* 2020;126(13):2971-9.
17. Duffy SW, Vulkan D, Cuckle H, et al. Effect of mammographic screening from age 40 years on breast cancer mortality (UK Age trial): final results of a randomised, controlled trial. *The Lancet Oncology.* 2020;21(9):1165-72.
18. Duffy S, Vulkan D, Cuckle H, et al. Annual mammographic screening to reduce breast cancer mortality in women from age 40 years: long-term follow-up of the UK Age RCT. *Health Technol Assess.* 2020;24(55):1-24.

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20. de Munck L, Siesling S, Fracheboud J, et al. Impact of mammographic screening and advanced cancer definition on the percentage of advanced-stage cancers in a steady-state breast screening programme in the Netherlands. *Br J Cancer*. 2020;123(7):1191-7.



# Breast Cancer: Age Specific New Cancer Cases

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## Mammography-Screening Potential Benefit and Harm

### Data background: Breast Cancer Surveillance Consortium Registry Data per 10.000 Women screened over 10 years

Age	40-49	50-59	60-69	70-74
Breast cancer death avoided (95% CI)	3 (0-9)	8 (2-17)	21 (11-32)	13 (0-32)
False-positive (n)	1212	932	808	696
Breast biopsies (n)	164	159	165	175
False-negative (n)	10	11	12	13
<b>Calculated from the data above</b>				
False-pos. to be accepted to avoid one breast cancer related death over 10 yrs per 10.000 women	404	117	38	54
Breast biopsies to be accepted to avoid one breast cancer related death over 10 yrs per 10.000 women	55	20	8	13

Mod. acc. Siu AL on behalf of the USPSTF 2016, 164:279–296

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2. Sicsic J, Pelletier-Fleury N, Moumjid N. Women's Benefits and Harms Trade-Offs in Breast Cancer Screening: Results from a Discrete-Choice Experiment. Value Health. 2018 Jan;21(1):78-88.



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## SSK Recommendation Moderate Risk

**Women with moderate risk increase from the age of 40: Moderate risk defined as a personal risk of disease of 15% to 29% (depending, for example, on the number and age of diseased family members, histologically proven risk lesions)**

Imaging modality	Grade of recommendation	Comment
Mammography	Primary examination	Individually adapted approach (depending on individual risk analysis, taking into account the woman's preferences and objections); in the age group of 50-69 years: Consider participation in the early detection program (mammography screening) and complementary procedures after benefit-risk analysis
US	Primary examination	Consider annually (especially in the case of high breast density)
Breast MRI	Special procedures	Only consider if a malignant finding is not sufficiently to exclude with mammography and sonography. There is insufficient data available for the use of MRI as a primary early detection method in the intermediate risk group

### Moderates Risiko berechnet nach Tyrer-Cuzick (QR-code)



Einfluß auf das Risiko haben auch z. B.: Alter, BMI, Brustdichte, frühere Brustbiopsien (Histologie), Vorerkrankungen (Ovarialkarzinom), Hormontherapie, Vorfahren (Ashkenazi Jewish), Alter bei Menarche, Alter bei erster Geburt.

[www.ssk.de/SharedDocs/Beratungsergebnisse/DE/2019/2019-06-27Orientie.html](http://www.ssk.de/SharedDocs/Beratungsergebnisse/DE/2019/2019-06-27Orientie.html)

Download 02. Januar 2025

## Früherkennung bei asymptomatischen Frauen Tomosynthese, Endpunkt: Cancer Detection Rate

	Oxford		
	LOE	GR	AGO
▪ Digitale Tomosynthese (DBT ± SM)*	1a	A	+
▪ Ersatz der DM durch synthetische MG + DBT**	1a	A	++

Es muss immer auch der komplette Datensatz der Tomosyntheseschichten zur Beurteilung zur Verfügung stehen, die alleinige synthetische Mammographie ist nicht ausreichend.

\* Bisher keine signifikante Reduktion der Intervallkarzinome.

Signifikant höhere Sensitivität, heterogene Spezifität und höhere Kosten [Gerät, Befunder, Archivierung] der digitalen Brust-Tomosynthese (DBT) im Vgl. zur digitalen Mammographie (DM).


\*\* Dosisreduktion durch Berechnung einer synthetischen Mammographie (SM) aus den Daten der DBT statt additiver DM.

1. Expert Panel on Breast Imaging; Niell BL, Jochelson MS et al. ACR Appropriateness Criteria® Female Breast Cancer Screening: 2023 Update. J Am Coll Radiol. 2024 Jun;21(6S):S126-S143.
2. Ciatto S, Houssami N, Bernardi D, et al.: Integration of 3D digital mammography with tomosynthesis for population breast-cancer screening (STORM): a prospective comparison study. Lancet Oncol 2017; 14 (7): 583-9, 2013
3. Houssami N, Bernardi D, Pellegrini M, et al. Breast cancer detection using single-reading of breast tomosynthesis (3D-mammography) compared to double-reading of 2D-mammography: Evidence from a population-based trial.(Storm-2) Cancer Epidemiol. 2017 Apr;47:94-99. doi: 10.1016/j.canep.2017.01.008.
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6. Hofvind S, Hovda T, Holen ÅS et al. Digital Breast Tomosynthesis and Synthetic 2D Mammography versus Digital Mammography: Evaluation in a Population-based Screening Program. Radiology. 2018 Jun;287(3):787-794. doi: 10.1148/radiol.2018171361. Epub 2018 Mar 1.
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- and mammography screening]. *Radiologe*. 2019 Jan;59(1):13-18. doi: 10.1007/s00117-018-0473-6. Review. German.
8. Marinovich ML, Hunter KE, Macaskill P et al. Breast Cancer Screening Using Tomosynthesis or Mammography: A Meta-analysis of Cancer Detection and Recall. *J Natl Cancer Inst*. 2018 Sep 1;110(9):942-949. doi: 10.1093/jnci/djy121.
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  10. Weigel S, Gerss J, Hense HW et al.: Digital breast tomosynthesis plus synthesised images versus standard full-field digital mammography in population-based screening (TOSYMA): protocol of a randomised controlled trial. *BMJ Open*. 2018 May 14;8(5):e020475. doi: 10.1136/bmjopen-2017-020475.
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  12. Kleinknecht JH, Ciurea AI, Ciortea CA. Pros and cons for breast cancer screening with tomosynthesis - a review of the literature. *Med Pharm Rep*. 2020;93(4):335-41.
  13. Giampietro RR, Cabral MVG, Lima SAM, et al. Accuracy and Effectiveness of Mammography versus Mammography and Tomosynthesis for Population-Based Breast Cancer Screening: A Systematic Review and Meta-Analysis. *Sci Rep*. 2020;10(1):7991.
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# Digital Breast Tomosynthesis

## European Guidelines on Breast Cancer Screening and Diagnosis

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 **Screening with tomosynthesis vs. mammography**  
*Issued on: May 2023*

**Cancer Screening, Diagnosis and Care**

**Healthcare question** **European Commission Initiative on Breast Cancer (ECIBC)**

Should screening using digital breast tomosynthesis vs. digital mammography be used in organised screening programmes for early detection of breast cancer in asymptomatic women?

**Recommendation**

**For asymptomatic women with an average risk of breast cancer, the ECIBC's Guidelines Development Group (GDG) suggests using digital breast tomosynthesis (DBT) over digital mammography (DM) in the context of an organised population-based screening programme.**

<https://cancer-screening-and-care.jrc.ec.europa.eu/en/ecibc/european-breast-cancer-guidelines?topic=65&usertype=60&updatef2=0>. Download am 09.01.2025

1. <https://cancer-screening-and-care.jrc.ec.europa.eu/en/ecibc/european-breast-cancer-guidelines?topic=65&usertype=60&updatef2=0>. Download am 09.01.2025

## KI zur Detektion in der Mammadiagnostik

	Oxford		
	LOE	GR	AGO
<b>KI im Screening</b>			
Als Second Reader in der Mammographie	<b>1b</b>	<b>B</b>	<b>+/-</b>
Zur Workload-Reduktion (AI alleine)	<b>2b</b>	<b>B</b>	<b>-</b>
Stand alone oder als Second Reader in der Tomosynthese	<b>2a</b>	<b>B</b>	<b>-</b>
Einsatz von KI noch zu früh, Überlegenheit und Übertragbarkeit in die klinische Routine nicht bewiesen, Standardisierung fehlt. [Uwimana A et al. 2025, Al-Karawi D et al. 2024]			

1. Uwimana A, Gnecco G, Riccaboni M. (2025) Artificial intelligence for breast cancer detection and its health technology assessment: A scoping review. *Comput Biol Med.* 2025 Jan;184:109391.
2. Al-Karawi D, Al-Zaidi S, Helael KA et al. Review of Artificial Intelligence in Breast Imaging. *Tomography.* 2024 May 9;10(5):705-726.
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6. Lång, K., V. Josefsson, A. M. Larsson, et al (2023). Artificial intelligence-supported screen reading versus standard double reading in the Mammography Screening with Artificial Intelligence trial (MASAI): a clinical safety analysis of a randomised, controlled, non-inferiority, single-blinded, screening accuracy study. Stand-Alone Artificial Intelligence for Breast Cancer Detection in Mammography: Comparison With 101 Radiologists. *Lancet Oncol.* England
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Digital Breast Tomosynthesis: A Systematic Review and Meta-Analysis." Radiology 307(5): e222639.

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Workload-Reduction:

1. Raya-Povedano, J. L., S. Romero-Martín, E. Elías-Cabot, et al (2021). "AI-based Strategies to Reduce Workload in Breast Cancer Screening with Mammography and Tomosynthesis: A Retrospective Evaluation." Radiology 300(1): 57-65.
2. Lång, K., S. Hofvind, A. Rodríguez-Ruiz and I. Andersson (2021). "Can artificial intelligence reduce the interval cancer rate in mammography screening?" Eur Radiol 31(8): 5940-5947.



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## Artificial Intelligence for Breast Cancer Detection and its Health Technology Assessment

	ML %			ML-unaided radiologist			ML-aided radiologists		
	Median	95%-C.I.		Median	95%-C.I.		Median	95%-C.I.	
<b>Sensitivity [%]</b>	<b>83.6</b>	79.7	86.9	<b>82.2</b>	77.2	86.3	<b>86.2</b>	80.6	90.3
<b>Specificity [%]</b>	<b>88.5</b>	84.0	91.9	<b>86.0</b>	81.9	89.2	<b>77.8</b>	57.1	90.2

While AI systems demonstrated promising applications across diverse breast imaging techniques, as highlighted in the reviewed articles and in our analysis indicating their superior diagnostic and predictive capabilities compared to conventional radiology practices, **our study underscores persistent challenges in developing and validating AI systems for clinical implementation**. These challenges stem from concerns regarding the availability and reliability of breast imaging data and the imperative to ensure the resilience, interpretability, and transparency of AI algorithms while navigating ethical and regulatory compliance considerations.

Uwimana A et al. *Comput Biol Med.* 2025 Jan;184:109391.

1. Uwimana A, Gnecco G, Riccaboni M. (2025) Artificial intelligence for breast cancer detection and its health technology assessment: A scoping review. *Comput Biol Med.* 2025 Jan;184:109391.

## Früherkennung (normales oder moderates Risiko\*) – Sonographie / MRT

	Oxford		
	LoE	GR	AGO
▪ <b>Screening-Mammasonographie alleine</b>	<b>5</b>	<b>D</b>	<b>--</b>
▪ <b>Autom. 3D-Sonographie</b>	<b>3a</b>	<b>C</b>	<b>-</b>
▪ <b>Mammasonographie als Ergänzung zur MG bei:</b>			
• <b>Dichtem Parenchym (inhomogen dicht, extrem dicht)</b>	<b>2a</b>	<b>B</b>	<b>++</b>
• <b>Erhöhtem Risiko</b>	<b>1b</b>	<b>C</b>	<b>++</b>
▪ <b>MRT bei neg. MG und extrem dichter Brust** 45-75 LJ</b>	<b>1b</b>	<b>B</b>	<b>+</b>

\* Außer INFP (intensiviertes Früherkennungsprogramm).

\*\* Definition von extrem dicht entspricht BIRADS-Dichtekategorie D, inhomogen dicht entspricht Kategorie C nach ACR BI-RADS Atlas 5. ed. 2013.

### ACR Expert Panel

1. Expert Panel on Breast Imaging; Niell BL, Jochelson MS et al. ACR Appropriateness Criteria® Female Breast Cancer Screening: 2023 Update. J Am Coll Radiol. 2024 Jun;21(6S):S126-S143.

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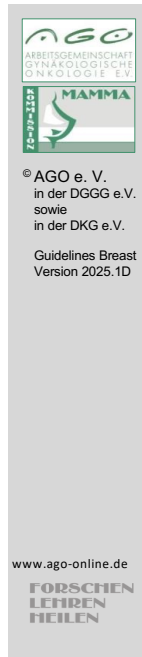
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## Früherkennung Klinische Untersuchung (Clinical Breast Examination; CBE)

	Oxford		
	LoE	GR	AGO
<b>Als alleinige Untersuchung</b>			
▪ <b>Selbstuntersuchung (BSE)</b>	1a	B	-*
▪ <b>Klinische Brust-Untersuchung (CBE) außerhalb der Krebsfrüherkennungsuntersuchung (KFU)</b>	1a	C	-*
▪ <b>Klinische Brust-Untersuchung (CBE) (im Rahmen der KFU)**</b>	1a	B	++
▪ <b>Medizinisch-taktile Untersuchung durch Blinde / Sehbehinderte</b>	3b	C	-
<b>CBE wegen klinisch- / mammo- / sonographischer Läsion</b>	5	D	++
<b>CBE in Kombination mit Bildgebung</b>	1a	A	++

\* Kann Brust-Bewusstsein erhöhen.

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## Abklärung von auffälliger Bildgebung (Screening oder diagnostisch) oder von Symptomen

	Oxford		
	LoE	GR	AGO
▪ <b>Klinische Untersuchung</b>	<b>3b</b>	<b>B</b>	<b>++</b>
▪ <b>Röntgenbasierte Bildgebung</b>			
▪ Mammographie (wenn noch nicht vorliegend)	1b	A	++
▪ Tomosynthese*	2a	B	+
▪ Kontrastmittelmammographie (CEM)	1b	B	+
▪ <b>Sonographie (B-mode)</b>	<b>2a</b>	<b>B</b>	<b>++</b>
▪ Elastographie (Shear wave)**	2b	B	+
▪ Automat. 3D-Sonographie	3b	B	+/-
▪ <b>Minimalinvasive Biopsie (CNB, VAB)</b>	<b>1b</b>	<b>A</b>	<b>++</b>
▪ <b>MRT***</b>	<b>2a</b>	<b>B</b>	<b>+</b>

\* Reduktion der Strahlenexposition durch DBT mit synthetischer Mammographie (SM) statt zusätzlich DM.

\*\* Zusatzuntersuchung

\*\*\* Wenn klinische, mammographische und sonographische Diagnostik inkl. CNB/VAB keine sichere Einschätzung erlauben.

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# CEM vs MRI

## European Guidelines on Breast Cancer Screening and Diagnosis

**Contrast-enhanced mammography**  
*Issued on: October 2021*



European  
Commission

Cancer Screening, Diagnosis and Care

**European Commission Initiative on Breast Cancer (ECIBC)**

**Healthcare question**


Should contrast-enhanced mammography vs. magnetic resonance imaging be used as additional imaging method to assist in surgical treatment planning in women with histologically confirmed invasive breast cancer?

**Recommendation**

In women with histologically confirmed invasive breast cancer, who require further evaluation, the ECIBC's Guidelines Development Group (GDG) suggests using contrast-enhanced mammography (CEM) over magnetic resonance imaging (MRI) as additional imaging method to assist in surgical treatment planning.

<https://cancer-screening-and-care.jrc.ec.europa.eu/en/ecibc/european-breast-cancer-guidelines?topic=65&usertype=60&updatef2=0>. Download am 09.01.2025

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## Comparison of Different Assessment Modalities

**Summary of pooled performance (95% confidence interval) of imaging modalities for recalled lesions.**

Imaging Modality	Sensitivity	Specificity
Digital mammography (DM)	85 (78-90)	77 (66-85)
Digital Breast Tomosynthesis (DBT)	91 (87-94)	85 (75-91)
Handheld Ultrasound (HHUS)	90 (86-93)	65 (46-80)
Contrast-enhanced Mammography (CEM)	95 (90-97)	73 (63-81)
Magnetic Resonance Imaging (MRI)	93 (88-96)	69 (55-81)

**Conclusions:** The CEM, MRI, DBT, and HHUS demonstrate excellent performance in correctly identifying and classifying cancer lesions referred for diagnostic work-up, but HHUS, MRI, and CEM have a more limited ability to discriminate benign lesions than DBT and DM.

*Akwo Cancers (Basel). 2024 Oct 17;16(20):3505. 2024*

Contrast enhancement and its pattern in CEM and MRI can be used to determine malignant or benign lesions. As some benign lesions show enhancement on CEM and MRI or hypervascularity on ultrasound, which increase the potential for FP; this may explain the lower overall specificity reported for HHUS, MRI, and CEM. Therefore, enhancement in MRI and CEM, and vascularity in ultrasound can be strengths and drawbacks at the same time.

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## Prätherapeutische Mammadiagnostik (nach histologischer Sicherung der Indexläsion)

	Oxford		
	LoE	GR	AGO
▪ Klinische Untersuchung	5	D	++
▪ Sonographie (Mamma)	2b	B	++
▪ Mammographie (wenn noch nicht vorliegend)	2b	B	++
▪ MRT*	1b	A	+
▪ Kontrastmittelmammographie (alleine) nach Verfügbarkeit und Strahlensensibilität der Brust (Alter)*	2a	B	+
▪ Tomosynthese + SM**	2b	B	+
▪ Mamma-CT	4	D	-
▪ Minimalinvasive Biopsie weiterer Mammabefunde (CNB, VAB)	1b	A	++
▪ Markierung des Tumors, wenn neoadjuvante Therapie geplant	1c	A	++

\* Möglichkeit der MRT-gestützten bzw. CEM-gestützter Biopsie (*in domo* oder im Rahmen eine Kooperation). MRT erwägen bei hohem familiärem Risiko, eingeschränkter Beurteilbarkeit in MG & US (Beurteilbarkeit C/D), invasiv lobulärem Karzinom, und ggf. vor neoadjuvanter Therapie.

\*\* Reduktion der Strahlenexposition durch DBT mit synthetischer Mammographie (SM) statt zusätzlicher DM

### DM, DBT, US, MRI

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## Prätherapeutische Axilladiagnostik

	Oxford		
	LoE	GR	AGO
▪ <b>Klinische Untersuchung</b>	5	D	++
▪ <b>Sonographie</b>	2a	B	++
▪ <b>CNB Axilla, wenn auffälliger LK-Befund und Markierung des LK wenn TAD geplant/≤ 3 susp. LK*</b>	2b	B	++
▪ <b>MRT</b>	1b	A	+
▪ <b>Mammographie (DM)</b>	2b	B	-
▪ <b>Tomosynthese (DBT)</b>	2b	B	-
▪ <b>Kontrastmittelmammographie (CEM, alleine)</b>	2a	B	-
▪ <b>PET für die Axilla (PET-CT, PET-MRT)</b>	2b	B	-
▪ <b>Mamma-CT</b>	4	D	-

\* Studienteilnahme empfohlen (AXSANA/EuBreast 3-Studie).

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### CEUS for Sentinel

#### Diagnosis of malignant lymph node infiltration

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#### Identification and tagging of the SLN (e.g carbon nanoparticle or wire guided)

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## Prätherapeutisches Staging

	Oxford		
	LoE	GR	AGO
▪ Anamnese und klinische Untersuchung	5	D	++
<b><u>Nur bei hohem Risiko für Fernmetastasen und/oder Symptomen und/oder Indikation zur (neo-)adjuvanten Chemo-/Antikörpertherapie:</u></b>			
▪ CT Thorax / Abdomen / Becken	2a	B	++
▪ Skelettszintigraphie	2a	B	+
▪ Röntgen-Thorax	5	C	+/-
▪ Leberzonographie	5	D	+/-
▪ Weiterführende Diagnostik je nach Befund (z. B. Leber-MRT / CEUS* / Biopsie etc.)	2a	B	+
▪ FDG-PET oder FDG-PET-CT** FDG-PET-MRT**	2a	B	+/-
▪ Ganzkörper MRT	2a	C	+/-

\* Contrast-enhanced ultrasound.
   
 \*\* Vorzugsweise bei hohem Stadium (III), wenn verfügbar.

### Statement: history and physical examination

1. GCP

### Statement: high metastatic potential / symptoms

1. Gerke O, Naghavi-Behzad M, Nygaard ST et al.. Diagnosing Bone Metastases in Breast Cancer: A Systematic Review and Network Meta-Analysis on Diagnostic Test Accuracy Studies of 2-[<sup>18</sup>F]FDG-PET/CT, <sup>18</sup>F-NaF-PET/CT, MRI, Contrast-Enhanced CT, and Bone Scintigraphy. Semin Nucl Med. 2025 Jan;55(1):137-151.
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## Diagnosing Bone Metastases in Breast Cancer: A Systematic Review and Network Meta-Analysis on Diagnostic Test Accuracy Studies of 2-[<sup>18</sup>F] FDG-PET/CT, <sup>18</sup>F-NaF-PET/CT, MRI, Contrast-Enhanced CT, and Bone Scintigraphy

Summary of Meta-Analyses. Estimates (and Respective 95% Confidence Intervals) of Sensitivity and Specificity Derived From Bivariate Random Effects Models and Network Meta-Analyzed Differences in Sensitivity and Specificity

Modality	Bivariate random effects models			Network meta-analysis <sup>o</sup>	
	Number of studies	Sensitivity	Specificity	Sensitivity	Specificity
2-[ <sup>18</sup> F]FDG-PET/CT	20	0.94 (0.89-0.97)	0.98 (0.96-0.99)	reference	reference
MRI	7	0.94 (0.82-0.98)	0.93 (0.87-0.96)	n.s.	n.s.
<sup>18</sup> F-NaF-PET/CT	4	0.95 (0.85-0.98)	1.00 (0.93-1.00)	n.s.	n.s.
CE-CT	5	0.70 (0.62-0.77)	0.98 (0.97-0.99)	p = 0.017	n.s.
Bone scintigraphy	13	0.83 (0.75-0.88)	0.96 (0.87-0.99)	p = 0.017	p = 0.053

<sup>o</sup> Network meta-analysis of multimodality studies (n = 16); comparator minus reference.

We concluded that 2-[<sup>18</sup>F]FDG-PET/CT and MRI have high and comparable accuracy for diagnosing bone metastases in breast cancer patients. Both outperformed CE-CT and bone scintigraphy regarding sensitivity. Future multimodality studies based on consented thresholds are warranted for further exploration, especially in terms of the potential role of <sup>18</sup>F-NaF-PET/CT in bone metastasis diagnosis in breast cancer.

*mod. from Gerke O et al. Semin Nucl Med. 2025 Jan;55(1):137-151.*

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## Neoadjuvante systemische Therapie Methoden zur Überprüfung des Ansprechens

	Oxford		
	LoE	GR	AGO
▪ Mammasonographie	2b	B	++
▪ Palpation	2b	B	++
▪ Mammographie	2b	B	++
▪ CEM (Kontrastmittel-verstärkte Mammographie)	2a	B	+
▪ MRT	2a	B	+
▪ DBT (Tomosynthese)	2b	B	+/-
▪ PET(-CT)	2b	B	+/-

Zur Kontrolle des Ansprechens sollte die gleiche Modalität prä- und posttherapeutisch gewählt werden.

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## Use of Different Imaging Modalities for Monitoring NACT

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**Comparison of HHUS, TOMO, MG and MRI: Murakami et al. AR Open 2021**

Size difference	<-11 mm	<-6 to -10mm	+/- 5 mm	>-6 to -10mm	>-11 mm
DBT vs. Patho	0%	5.3%	82.1%	8.4%	4.2%
FFDM vs Patho	9.5%	9.5%	62.1%	8.4%	10.5%
US vs Patho	10.5%	9.5%	61.1%	12.6%	6.3%
MRI vs. Patho	3.2%	8.4%	75.8%	6.3%	6.3%

**CEM compared to MRI: Kaiyin et al. BCRT 2023**

Six head-to-head comparison studies with 328 patients included.  
Pooled **sensitivity, specificity, and diagnostic odds ratio (DOR)**:

**CEM:** 93% (95% CI, 84-97%), 68% (CI, 60-76%), 29.29 (CI, 11.41-75.18)  
**MRI:** 84% (95% CI, 62-95%), 80% (CI, 71-87%), 21.39 (CI, 5.94-77.13).  
**AUC: CEM:** 0.85 (95% CI 0.82-0.88)    **MRI:** 85 (95% CI 0.82-0.88)

**CEM depending on Ca-subtype: Vidali et al. 2024**

**Conclusions:**  
Overall, CEM is accurate in assessing the pCR and predicting the pathologic-complete response among the different molecular subtypes after NAT.

**Conclusion:**  
DBT has good correlation with histopathology for measuring residual tumor size after NST. DBT was comparable to MRI in assessing tumor response after completion of NST.

**Conclusion:**  
This meta-analysis showed that CEM provides an equivalent diagnostic accuracy to MRI in identification of pCR in breast cancer patients receiving NAT. The results support the increasing use of CEM in this setting and would encourage future studies to validate CEM as a suitable replacement for MRI.

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