

Molecular Breast Imaging May Benefit Women with Dense Breasts

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OAK BROOK, Ill. – Screening women with dense breasts with both molecular breast imaging (MBI) and digital breast tomosynthesis (DBT) increased overall invasive cancer detection while modestly increasing the recall rate compared with screening only with DBT, according to a new study published today in *Radiology*, a journal of the Radiological Society of North America (RSNA).

"To our knowledge, this is the first multicenter, prospective evaluation of MBI as a supplement to DBT in women with dense breasts," said lead author Carrie B. Hruska, Ph.D., professor of medical physics at Mayo Clinic in Rochester, Minnesota.

An estimated 47% of women who undergo breast cancer screening have dense breasts, according to the Centers for Disease Control and Prevention. DBT is an advanced form of mammography that takes multiple X-ray images of the breast from different angles to create a 3D reconstruction of the breast, but it does not detect all breast cancers, especially in women with dense breasts.

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Carrie B. Hruska, Ph.D.

MBI is one of several options available for supplemental breast screening for dense breasts, such as breast ultrasound, breast MRI and contrast-enhanced mammography. The Density MATTERS (Molecular Breast Imaging and Tomosynthesis to Eliminate the Reservoir) Trial was designed to assess the performance of screening MBI as a supplement to DBT in women with dense breasts.

For the trial, women with dense breasts from five sites were prospectively enrolled from 2017 to 2022 and underwent two annual screening rounds of DBT and MBI (prevalence screening at Year 1 and incidence screening at Year 2). One-year follow-up after each screening round was completed in September 2024.

Eligible participants included women aged 40–75 years who were asymptomatic and had dense breasts as visually assessed by a radiologist and reported on their last mammogram.

The study cohort included 2,978 participants (mean age 56.8 years). The women were mostly postmenopausal, and 82% had category C breast density. Approximately 80% of participants had no family history of breast cancer, and 98% had no personal history of breast cancer.

Across both screening rounds, 30 breast cancer lesions were detected in 29 participants by MBI only and not found with DBT. Most of these incremental breast cancers were invasive (22 of 30 or 71% of lesions). The median invasive lesion size was 0.9 cm. Among the participants with MBI-only detected breast cancer, 26 of 29 (90%) had node-negative cancers, and 6 of 29 (20%) had node-positive disease.

"MBI detected an additional 6.7 cancers per 1,000 screenings at Year 1 and an additional 3.5 cancers per 1,000 screenings at Year 2," Dr. Hruska said. "Among the incremental cancers detected only by MBI, 70% were found to be invasive. Additionally, 20% of those incremental cancers were node positive, suggesting that MBI can reveal mammographically occult, clinically important disease."

In the first screening round, 7 of 2,978 participants (2.4 per 1,000 screened) were diagnosed with node-positive cancers. DBT alone detected 4 of 7 (57%), and DBT plus MBI detected 7 of 7 (100%).

In Year 2, 6 of 2,590 participants (2.3 per 1,000 screened) had node-positive cancers. DBT alone detected 1 of 6 (16%) and DBT plus MBI detected 4 of 6 (67%). Neither modality detected 2 of the 6 node-positive cancers (33%).

"Someone who's having their routine annual screen every year should not be diagnosed with advanced breast cancer," Dr. Hruska said. "That's just unacceptable. With a supplemental screening every few years, we hope to find cancers earlier and see the diagnosis of advanced cancer go way down."

Dr. Hruska said one of the strengths of the Density MATTERS trial was the mix of academic medical centers and community hospitals involved. Participating centers also included MD Anderson Cancer Center, Henry Ford Health System, ProMedica Breast Care (regional practice in Ohio), and a Mayo Clinic Health System site in La Crosse, Wisconsin.

"The enrollment of 12% minority patients extends the generalizability of our findings," she said.

She said the trial results provide important data for healthcare institutions assessing the best modality for supplemental breast screening.

"I don't want to discourage anyone from getting a mammogram, because they absolutely should," Dr. Hruska said. "However, DBT doesn't find all cancers, and women need to understand its limitations and consider how supplemental screening can fill the gap."

According to Dr. Hruska, MBI is considered safe for routine screening, is well-tolerated by patients and is relatively inexpensive.

"MBI uses a well-established radiotracer that's been used in cardiac imaging for a really long time," she said. "It has fewer risks than other modalities and no contrast reactions. If a woman has a choice of modalities, it's important that she understands the benefits and risks of each and be involved in the decision-

making."

"Molecular Breast Imaging and Digital Breast Tomosynthesis for Dense Breast Screening: The Density MATTERS Trial." Collaborating with Dr. Hruska were Katie N. Hunt, M.D., Nicholas B. Larson, Ph.D., Patricia A. Miller, M.D., Richard L. Ellis, M.D., Robin B. Shermis, M.D., Gaiane M. Rauch, M.D., Ph.D., Amy Lynn Connors, M.D., Jeannette Gasal Spilde, M.D., Dominic T. Semaan, M.D., Emily C. Siegal, M.D., Shannon N. Zingula, M.D., Sabala R. Mandava, M.D., Tamara S. Martin, M.D., Riflat K. Ahmed, M.D., Dana H. Whaley, M.D., Beatriz Adrada, M.D., Lacey R. Gray, C.N.M.T., Ramy A. Mehta, M.S., Rebecca J. Roll, Roberta E. Redfern, Ph.D., Michael O'Connor, Ph.D., and Deborah J. Rhodes, M.D.

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For patient-friendly information on breast imaging, visit [RadiologyInfo.org](https://radiologyinfo.org).

Images (JPG, TIF):

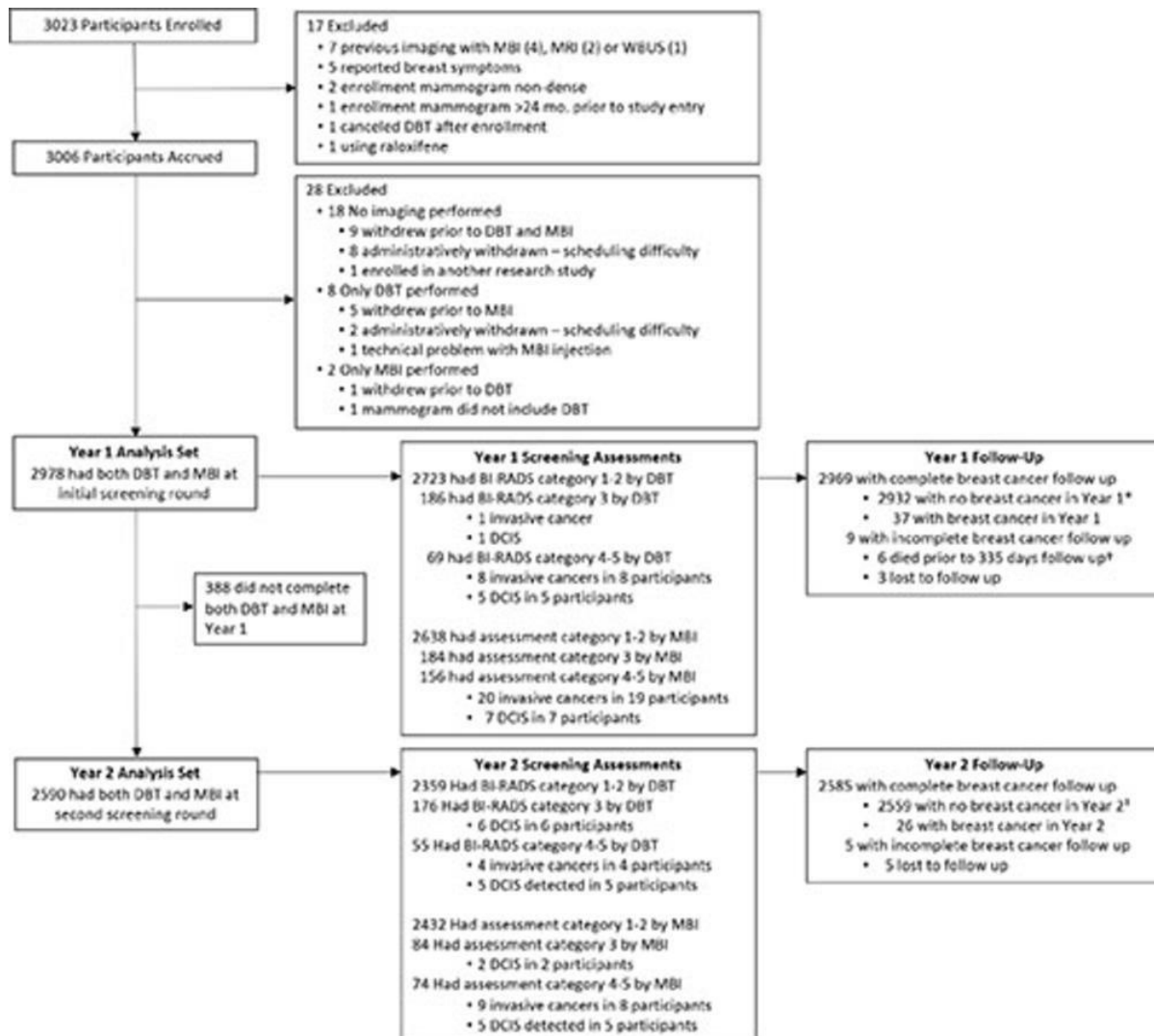


Figure 1. Participant flowchart. BI-RADS = Breast Imaging Reporting and Data System, DBT = digital breast tomosynthesis, DCIS = ductal carcinoma in situ, MBI = molecular breast imaging, WBUS = whole-breast US. * = Of the 2932 participants with no breast cancer in year 1, follow-up was completed with negative or benign breast imaging (n = 2769), chart review (n = 133), or participant contact (n = 30) at least 335 days after the year 1 screening. † = Deaths were not due to breast cancer. ‡ = Of the 2559 participants with no breast cancer in year 2, follow-up was completed with negative or benign breast imaging (n = 2165), chart review (n = 308), or participant contact (n = 86) at least 335 days after the year 2 screening.

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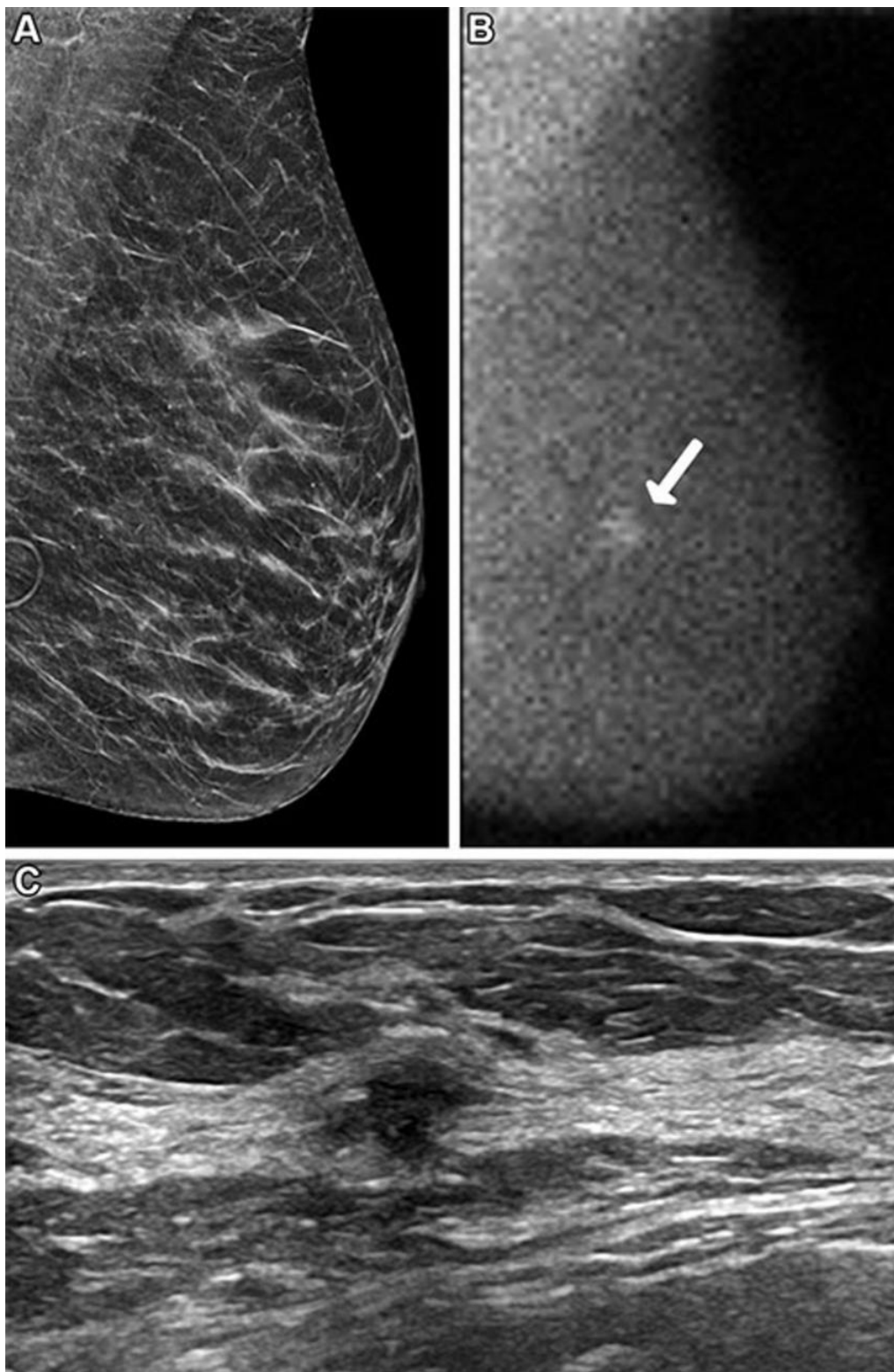


Figure 2. Images in a 62-year-old woman who presented for screening. **(A)** Image from digital breast tomosynthesis screening (the synthesized two-dimensional mediolateral oblique view of the left breast is shown) at year 1 was interpreted as negative and showing heterogeneously dense breast. **(B)** Image from molecular breast imaging screening (mediolateral oblique view) at year 1 reveals a 0.9-cm nonmass focal area of uptake in the left breast (arrow). **(C)** Targeted US scan (transverse image) shows a suspicious mass in the 3-o'clock position of the left breast, 4 cm from the nipple. US-guided core biopsy and lumpectomy revealed a 0.9-cm grade 2 invasive ductal carcinoma, an estrogen receptor-negative, progesterone receptor-negative, human epidermal growth factor receptor 2-negative lesion, with two sentinel nodes negative (N0).

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Figure 3. Images in a 57-year-old woman who presented for screening. **(A)** Image from digital breast tomosynthesis screening (synthesized two-dimensional mediolateral oblique view of the left breast) at year 1 was interpreted as negative and showing heterogeneously dense breast. **(B)** Image from molecular breast imaging screening (mediolateral oblique view) at year 1 reveals a 2-cm mass in the left breast (arrow). **(C)** Targeted US scan (transverse image) reveals a corresponding irregular hypoechoic mass in the 1-o'clock position, 5 cm from the nipple. US-guided core biopsy and lumpectomy revealed a 2.6-cm grade 2 invasive ductal carcinoma, an estrogen receptor–positive, progesterone receptor–negative, human epidermal growth factor receptor 2–positive lesion, with two sentinel nodes negative (N0).

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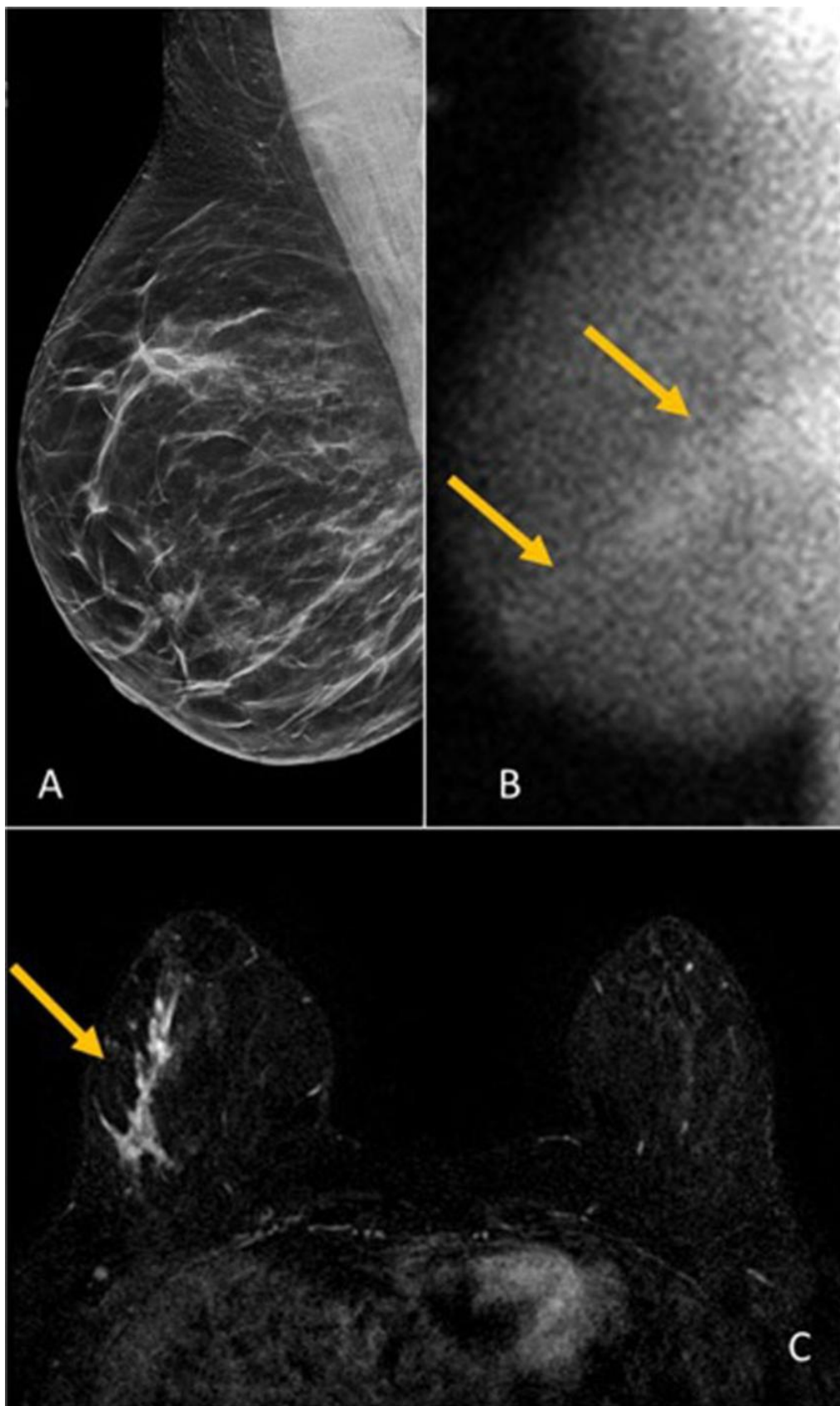


Figure 4. Images in a 48-year-old woman who presented for screening. **(A)** Image from digital breast tomosynthesis screening (synthesized two-dimensional mediolateral oblique view of the right breast) at year 1 was interpreted as negative and showing heterogeneously dense breast. **(B)** Image from molecular breast imaging screening (mediolateral oblique view) at year 1 reveals a nonmass segmental area of uptake in the right breast (arrows). **(C)** Contrast-enhanced MRI scan (subtraction maximum intensity projection) shows corresponding segmental nonmass enhancement measuring 9.8 cm to the greatest extent (arrow). MRI-guided core biopsy and lumpectomy revealed high-grade ductal carcinoma in situ, 2.5 cm in extent, estrogen receptor-negative, and progesterone receptor-negative lesions.

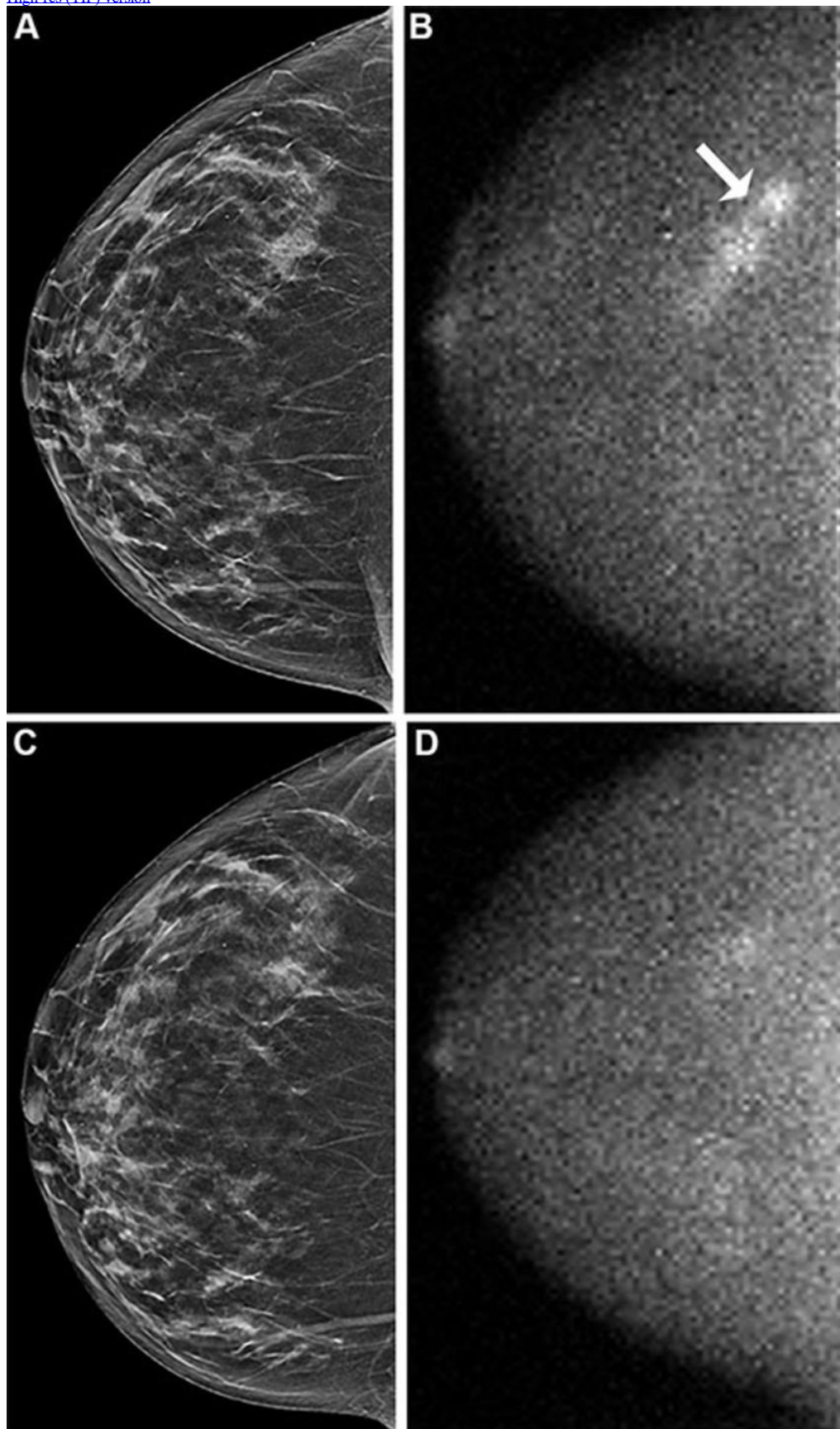


Figure 5. Images in a 69-year-old woman who presented for screening. **(A)** Image from digital breast tomosynthesis (DBT) screening (synthesized two-dimensional craniocaudal view of the right breast) at year 2 was negative and showing heterogeneously dense breast. **(B)** Molecular breast imaging (MBI) screening (right craniocaudal view) at year 2 shows a 4.7-cm nonmass focal area of uptake (arrow). US-guided biopsy and mastectomy revealed a grade 2 invasive lobular carcinoma, 9.1 cm in greatest extent, that was estrogen receptor positive, progesterone receptor positive, and human epidermal growth

factor receptor 2 negative, with two sentinel nodes and 10 axillary nodes negative (N0). **(C)** Prior year 1 DBT was interpreted as negative. **(D)** Prior year 1 MBI was interpreted as negative with bilateral moderate background parenchymal uptake; a retrospective review revealed that mild uptake correlated with the location of the lesion detected at year 2.

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