

Clinical Experience in Contrast-enhanced Mammography

By Andrea Woodroof, MD

Background

There have been many recent advancements in breast imaging. In particular, screening tomosynthesis has been shown to improve cancer detection while decreasing recall rates. The use of synthesized 2D images, calculated from the tomosynthesis datasets, allows similar improvements to cancer detection while lowering the radiation dose of 2D/3D combination imaging to levels of 2D mammography. The advent of tomosynthesis-guided biopsies allows us to completely evaluate lesions detected on these new technologies.

As impressive as these recent advancements are, their gains are mainly in the area of increasing the sensitivity and specificity of screening exams for the detection of breast cancer detection. As we know, functional imaging yielding data about the physiologic activity of a lesion (such as that given by dynamic contrast enhanced breast MRI or PET imaging) has historically been unavailable in the mammographic modality. This has been particularly frustrating, given that mammography is the only modality that frequently offers radiologists multiple prior examinations by which to establish stability of findings—which often obviates the need for further work up or biopsy of lesions.

However, things are changing. Contrast mammography, which uses standard iodinated contrast agents, can now be used in conjunction with mammography equipment to answer physiologic

questions about a breast lesion that breast MRI has historically answered. Published studies have shown contrast mammography to have equal or near equal sensitivity of MRI, but with higher specificity. In 2013, Hologic received clearance to market contrast-enhanced 2D mammography (CE2D) on its Selena® Dimensions® system. This system, which is primarily a software upgrade to the company's Dimensions unit, allows one to perform contrast procedures at the point of diagnostic evaluation, and does so with shorter procedure times, easier access, and lower costs as compared to breast MRI.

The procedure can be performed as a 2D contrast image or a 2D contrast image combined and co-registered to a tomosynthesis dataset. In both cases the procedure creates both a morphological image (standard 2D and/or tomosynthesis image) and the functional image (contrast 2D) for review. We perform both types of imaging in our practice, depending on the case.

About Our Practice

We are a small breast clinic located in Southern Kentucky that serves 8,000 patients per year with both screening and diagnostic imaging services. In our rural geographical area, MRI breast imaging has historically been challenging because there is no immediate access to high-quality breast MRI. When a facility in our area did upgrade to offer breast MRI, the facility did not offer MRI-guided biopsy, which meant that patients with MRI-detected BI-RADS 4 lesions were directed to an out-of-state facility to undergo repeat MRI and, if indicated

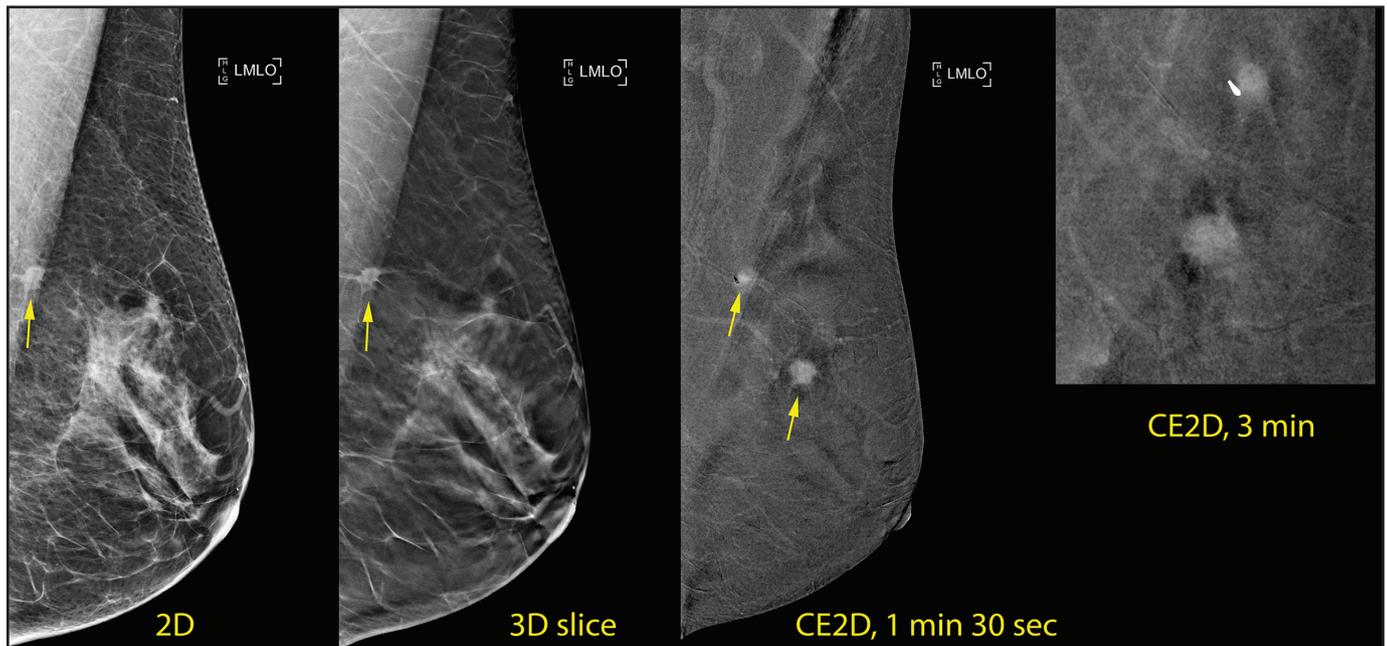
at that point, MRI-guided biopsy. We wanted to offer accessible, accurate, and affordable services to our patients in need of contrast-enhanced imaging, and we needed the capacity to biopsy suspicious lesions that we discovered.

Our challenge, like most practices, is providing value-based care. We began using CE2D in the September of 2014 in order to expand our range of services and offer better, cost effective care to patients. Since then, we have used CE2D in a variety of clinical applications that historically would have been evaluated with MRI. Specifically, we have used CE2D for the evaluation of patient-reported palpable regions of concern that were radiographically normal on diagnostic work-up, patients with pacemakers, patients with breast cancer to evaluate for extent of disease, radiology pathology discordance, claustrophobia, the uninsured, the underinsured and those with no access to transportation to outside facilities with MRI. It has been remarkably clinically effective to use one single modality to perform digital mammography, breast tomosynthesis, tomosynthesis-guided biopsy, and contrast imaging.

How It Works

The procedure can be broken down into two steps: contrast administration and imaging. The contrast agent is a standard non-ionic CT contrast agent that is delivered intravenously to a patient using a power injector. After waiting two minutes for the contrast agent to distribute into the breast, standard mammography imaging proceeds. One can image either or both breasts in any projection desired.

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CASE 1: Tomosynthesis reveals a spiculated mass in the left axillary tail and a suspicious 1 cm mass (discordant in U/S) within the dense band of tissue at the 2:00 axis of the left breast.

For a given image, the system takes two separate but nearly simultaneous exposures, the first image is a standard 2D mammogram and the second is a kV mammogram. These images are subtracted automatically, resulting in a CE2D image that highlights any focal areas of enhancement. If one desires, one can also take a tomosynthesis series as part of the contrast imaging protocol. These images are co-registered to the 2D FFDM and can be used for biopsy guidance using the Affirm® breast biopsy system of the same gantry.

The entire procedure takes less than 10 minutes and is broken out as follows in our experience:

- Injection of iodine. The patient is seated.
- After 2 minutes, we begin positioning the patient
- We have an approximately 6-minute window where the contrast agent can be well visualized. In that time we can easily perform CC and MLO images of both breasts, and if desired, additional projections to show enhancement to best advantage.

Use of CE2D in Clinical Practice

We find CE2D is useful for a variety of circumstances. Specifically, we have

used CE2M to evaluate discordant radiographic and pathologic findings.

CE2D for Discordant Findings

CASE 1: 52-year-old female who presented for screening, which demonstrated a 0.8 cm spiculated mass in the left axillary tail.

Screening 2D/Tomosynthesis showed

1. A 0.8 cm spiculated mass in the left axillary tail.
2. A 0.8 cm well circumscribed mass at the 2:00 axis 2 cm from the nipple seen on tomosynthesis. Adjacent to the mass, there is a focal area of possible distortion noted.

Ultrasound demonstrated a 0.6 cm irregular isoechoic mass at the 1:00 axis 9 cm from the nipple corresponding to the 0.8 cm spiculated mass in the left axillary tail seen mammographically. In addition, it showed a dense band of tissue along the 2:00 axis and within the band, a well circumscribed 0.6 cm avascular hypoechoic mass with some acoustic enhancement is seen. The distortion seen mammographically does not show a definite sonographic correlate and therefore cannot be biopsied under sonographic guidance.

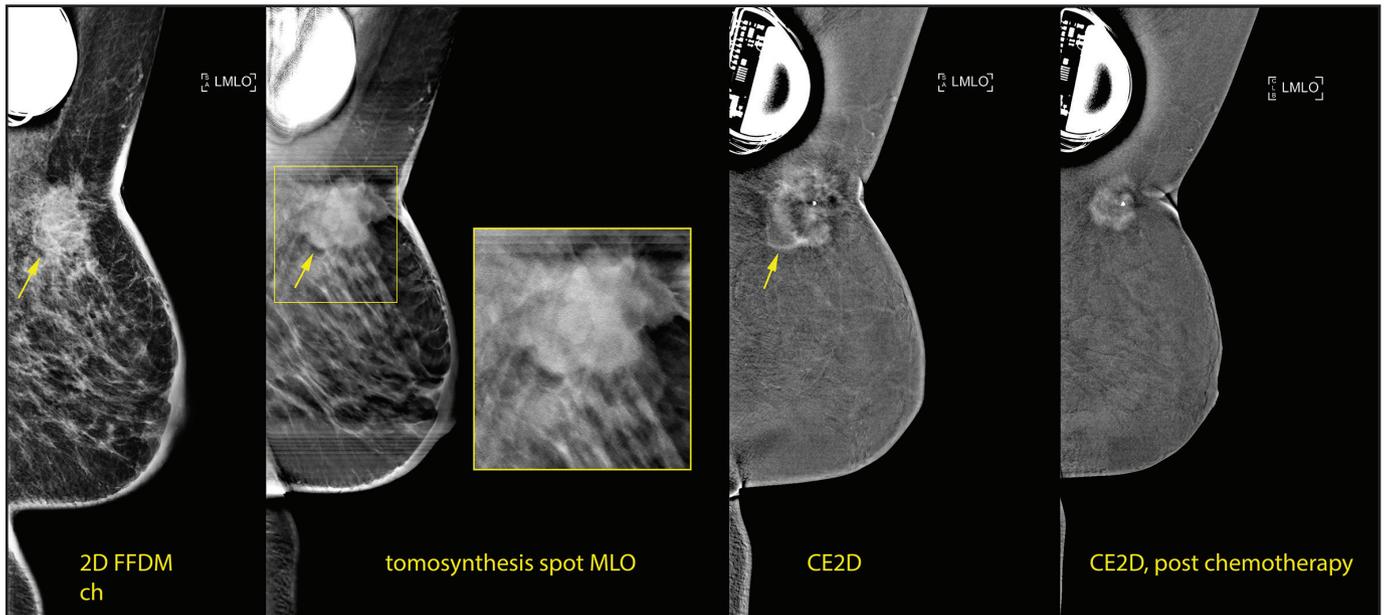
Ultrasound-guided biopsy of the spiculated lesion in the left axillary tail was

performed and post-procedural mammogram demonstrates accurate placement of the clip within the mass. However, pathology results were discordant, reporting only Fibroadipose Tissue. Stereotactic biopsy was planned for the discordant lesion in the axillary tail and the distortion seen on tomosynthesis.

Prior to tomosynthesis-guided biopsy, CE2D was performed showing suspicious intense contrast enhancement of the spiculated mass in the left axillary tail and suspicious enhancement of a 1 cm mass within the dense band of tissue at the 2:00 axis of the left breast.

The enhancing mass was cross-localized from the 2D image of the contrast exam to the 2D/Tomosynthesis exam, documenting that the distortion seen on tomosynthesis corresponded to the enhancing mass seen on CE2D. The ability to directly trace a focus of enhancement to its mammographic correlate without crossing modalities increases confidence that the etiology of enhancement is understood, and the stability of the mammographic correlate can then be directly ascertained.

Tomosynthesis-guided biopsy of the spiculated mass in the axillary tail and the distortion at the 2:00 axis of the left breast was performed. Pathology



CASE 2. Spiculated mass in the left breast with skin thickening and retraction as seen in the 2D, tomosynthesis slice and contrast enhanced mammography image in a MRI-contraindicated patient.

results of the spiculated lesion reported a columnar change and ductal epithelial hyperplasia. Pathology of the distortion reported a ruptured cyst with surrounding fibrosis with sclerosing adenosis with no evidence of atypia or carcinoma.

Radiographically, these were discordant, so surgical excision of both areas was performed. Surgical excision pathology demonstrated invasive carcinoma with tubular features for both lesions.

CE2D for MRI-contraindicated Patients

We have also found CE2D useful for patients in which MRI is not an option, such as those with generous body habitus, pacemaker placements, and those not amenable to MRI secondary to claustrophobia, lack of insurance coverage, or high deductible costs.

CE2D for Pre- and Post-neoadjuvant Chemotherapy in a Patient With Pacemaker

CASE 2: 50 year-old female, presenting with a six month history of a “pin-sized area” of irritation underlying the skin of her left breast.

The patient assumed the irritation was related to her pacemaker and waited for the irritation to resolve. Over 6 months, the pin-sized area of irritation developed into a 4 cm erythematous and

indurated mass-like thickening overlying her pacemaker.

Diagnostic evaluation with 2D/tomosynthesis demonstrated a 4 cm spiculated mass at the 1:00 axis of the left breast with associated skin thickening and skin retraction. Ultrasound showed a 3.5 cm solid irregular mass at the 1:00 axis, with the left axilla containing abnormal lymph nodes lacking fatty hila.

Ultrasound-guided biopsy of the spiculated mass and of one axillary lymph node was performed. Pathology results showed invasive ductal carcinoma within the breast mass; and metastatic ductal carcinoma involving the biopsied axillary lymph node. The patient was referred to oncology for neoadjuvant chemotherapy.

Because the transcutaneous pacemaker negated the option of pre-treatment and post-treatment MRI to evaluate response to treatment, and PET was declined by insurance, CE2D was performed.

CE2D prior to neoadjuvant therapy showed strong enhancement of the mass at the 1:00 axis of the left breast. The 3.4 × 3.4 cm mass can only be seen on the MLO view secondary to fixation to the chest wall.

Following neoadjuvant chemotherapy, the CE2D exam shows strong enhancement of the mass at the 1:00 axis of the left breast, with an interval decrease

in the size of the mass, now measuring 1.7 × 1.7 cm. Interval decrease in skin thickening of the right breast also seen and the mass can now be pulled into view on XCC imaging.

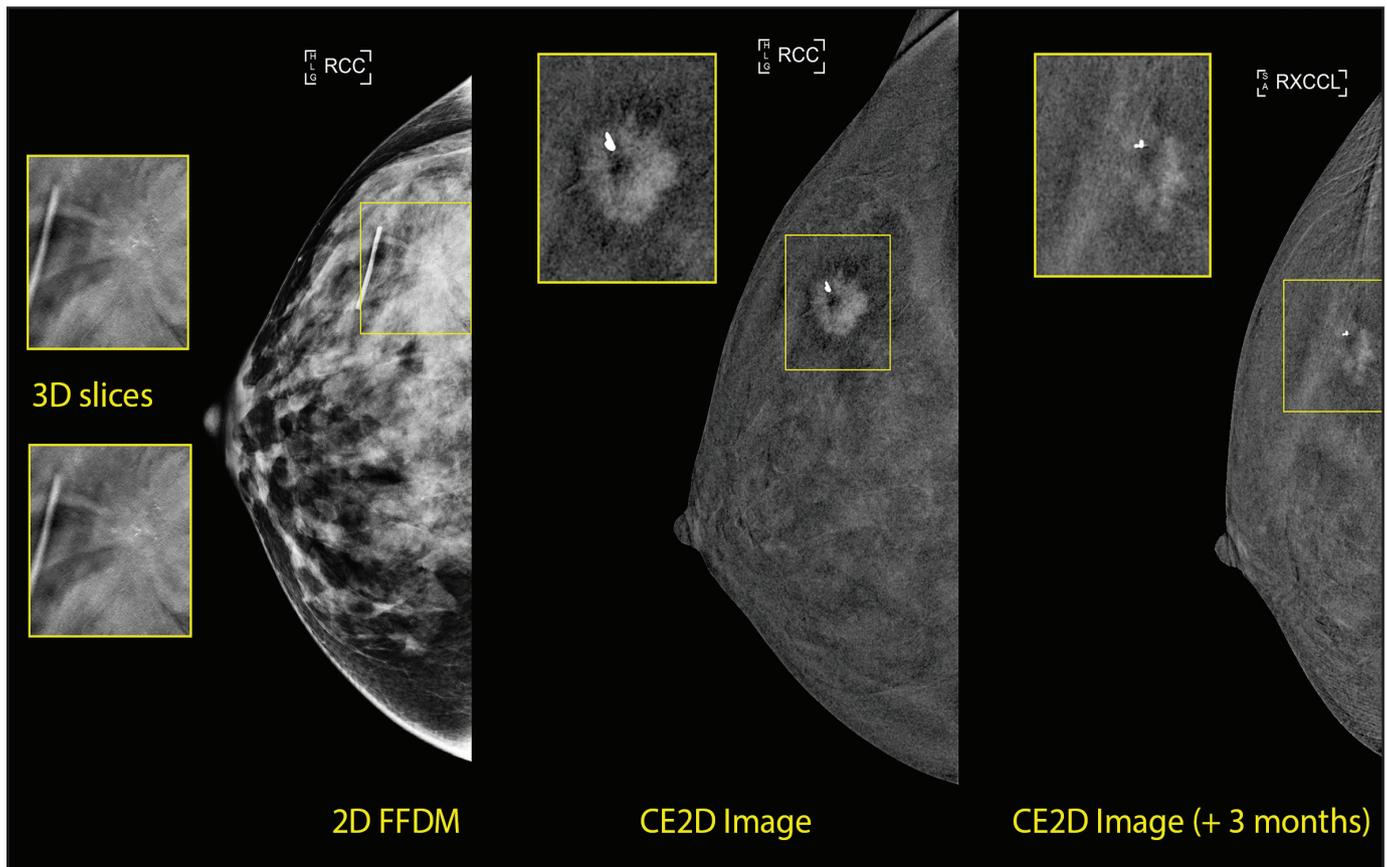
CE2D for Measuring Extent of Disease in Very Dense Breasts and for Evaluating Response to Neoadjuvant Chemotherapy

CASE 3: 49 year-old female presenting with a palpable mass with skin dimpling in the right breast

This patient underwent diagnostic 2D/tomosynthesis which revealed a spiculated mass with associated calcifications at the 11:00 axis of the right breast. Prominent lymph nodes were also noted in the axilla. Because the mass was embedded within very dense parenchymal tissue, the extent of disease was not easily measured by standard mammography.

Ultrasound demonstrated a 1.5 cm hypoechoic mass at the 11:00 axis 4 cm from the nipple corresponding to the spiculated mass seen mammographically. Ultrasound-guided biopsy of the mass and biopsy of a prominent lymph node was performed. A clip was placed within the mass, and follow-up mammography demonstrated accurate clip placement.

Pathology results demonstrated invasive ductal carcinoma within the



CASE 3. Spiculated mass seen in mammography and further localized with contrast enhanced mammography. The contrast-enhanced 2D mammogram imaged three months later reveals a partial response to therapy.

mass, as well as metastatic invasive ductal carcinoma involving the biopsied lymph node. The patient was referred for oncological evaluation and neoadjuvant chemotherapy.

Prior to initiation of neoadjuvant chemotherapy, CE2D was performed, confirming the presence of a 1.5 cm × 1.3 cm enhancing spiculated mass at the 11:00 axis of the right breast.

After the of neoadjuvant therapy, repeat CE2D was performed to evaluate response. The enhancing mass showed an interval decrease in size, demonstrating a partial response to neoadjuvant chemotherapy.

The patient then underwent bilateral mastectomy with right axillary lymph

node dissection. Surgical pathology demonstrated the residual tumor within the upper outer quadrant of the right breast, measuring a maximum of 1 cm.

Conclusion

We have found contrast mammography to be an excellent adjunct to our practice. In our preliminary experience, CE2D has been a clinically effective, efficient, and inexpensive tool for evaluating breast tissue and breast cancer. The advancements in this technology have allowed us to utilize one machine for screening, diagnostic evaluation, biopsy, and response to treatment. The cost effectiveness of this technology has been an excellent fit for our practice.

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