There should be no delay in screening mammograms after COVID-19 vaccination due to a swelling in the lymph nodes called lymphadenopathy, according to a new study published in the journal *Radiology*. Instead, lymphadenopathy should be interpreted within the context of the individual patient's risk factors, the researchers said.

Lymph nodes commonly become enlarged in response to an infection or vaccine, but unilateral (affecting one side of the body) lymphadenopathy that occurs suddenly may also signal cancer. Therefore, unilateral lymphadenopathy, a known side effect of the COVID-19 vaccines, has posed a diagnostic challenge in oncologic imaging.

When this side effect was first discovered, women were told to delay their mammography exams to avoid unnecessary false positives and follow-ups. However, it soon became apparent that the swelling could take many weeks to resolve.

Therefore, Stacey Wolfson, M.D., from the Department of Radiology at New York University Grossman School of Medicine and NYU Langone Health in New York City, and colleagues set out to determine the outcomes of axillary lymphadenopathy (swelling of the lymph nodes in the armpits) after COVID-19 vaccination on breast imaging examinations.
"This is the largest study to evaluate axillary lymphadenopathy after COVID-19 vaccination, with long term follow-up data," Dr. Wolfson said. "This is important because it gives us further insight as to the natural behavior of this reactive lymphadenopathy and allows us to make broader guidelines regarding follow-up."

The researchers identified patients who received the COVID-19 vaccine and had breast imaging between December 30, 2020, and April 12, 2021, at 17 sites across one institution. Follow-up exams were recorded through December 10, 2021. The research team analyzed clinical data and vaccine information from the medical records, and only patients with complete information were included.

Of the 1,217 patients who received the COVID-19 vaccination and had breast imaging, 537 (44%) had lymphadenopathy identified on at least one imaging exam, 823 patients (68%) had screening exams, and 334 patients had mammography and sonography performed the same day. Of these, 29 (9%) patients had lymphadenopathy identified on mammography alone, 203 (61%) patients on ultrasound alone, and 102 (30%) patients on both exams. In patients with lymphadenopathy, the average node measured 1.8 centimeters.

"Other studies with lower reported incidence of lymphadenopathy are based on self-reported symptoms or detection solely on mammography. In contrast, our study had an incidence of 44%," Dr. Wolfson said.

Of the patients with known vaccine manufacturer information, 459 patients received the Moderna vaccine (46% with lymphadenopathy), 505 patients received the Pfizer vaccine (38% with lymphadenopathy), and 18 patients received the Johnson & Johnson vaccine (39% with lymphadenopathy).

Patients developed lymphadenopathy as early as one day following the first vaccine dose and as late as 71 days following the second dose. Persistent axillary lymphadenopathy was seen up to 43 weeks after vaccination.

"I was surprised by how quickly the lymph nodes became swollen and how long they
persisted after being detected on routine screening mammogram and screening ultrasound exams," Dr. Wolfson said. "We found benign reactive lymph nodes were still present despite delaying the screening exams for 4 to 6 weeks based on various guidelines. These lymph nodes were unchanged with follow up exams at three months, and some enlarged lymph nodes persisted for over 10 months."

Eight percent (43/537) of patients with lymphadenopathy underwent a biopsy. Among them, 34 (79%) had benign results and 9 (21%) had malignant results. Four patients were diagnosed with metastatic breast cancer. The patients with metastatic cancer all had suspicious concurrent mammographic findings in the ipsilateral breast.

Four patients were diagnosed with lymphoma. Three of these patients already had known diagnoses, and the fourth patient had bilateral lymphadenopathy. One patient with a known history of lung cancer was diagnosed with lung cancer metastatic to an axillary lymph node.

Three hundred eighty-seven (72%) patients with lymphadenopathy had 407 follow-up examinations, averaging 15.7 weeks after the initial examinations. On follow-up imaging, 323 of 407 (79.4%) exams were assessed as benign and 84 (20.6%) were assessed as probably benign with recommendation to undergo additional follow-up. No patients in the follow-up group were diagnosed with a subsequent malignancy.

"These findings allow us to make broader guidelines suggesting that short-term follow-up imaging is not recommended," Dr. Wolfson said. "We found only 9 cancerous lymph nodes in our study, and they were only found in women with known cancer, concurrent suspicious findings in the breast, and/or contralateral axilla."

Based on the findings, the researchers concluded that there should be no delay in screening mammograms because of recent COVID-19 vaccination and lymphadenopathy should be interpreted in the context of patient risk factors with vigilance in patients with concurrent suspicious mammographic findings in the ipsilateral breast.

"It is important for people to know that they should not delay their screening mammograms due to recent vaccination. Reactive lymphadenopathy is common after COVID-19 vaccination – and benign," Dr. Wolfson said.

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"Axillary Adenopathy after COVID-19 Vaccine: No Reason to Delay Screening Mammogram." Collaborating with Dr. Wolfson were Eric Kim, M.D., Anastasia Plaunova, M.D., Rita Bukhman, M.D., Ruth D. Sarmiento, M.D., Naziya Samreen, M.D., Divya Awal, M.D., Monica M. Sheth, M.D., Hildegard B. Toth, M.D., Linda Moy, M.D., and Beatriu Reig, M.D., M.P.H.

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