Regression in local recurrence in the contralateral breast following mastectomy in bilateral locally advanced breast cancer: A comparison of neutrophil-to-lymphocyte and platelet-to-lymphocyte ratios

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INTRODUCTION

Breast cancer is the most commonly observed type of cancer in women; in approximately 25%-30% of patients, diagnosis is made when the breast cancer is locally advanced (1). As in other cancers, the inflammatory response in breast cancer plays an important role in cancer development. The neutrophil-to-lymphocyte ratio (NLR) and platelet-to-lymphocyte ratio (PLR), which can be obtained from a peripheral full blood count, are routinely clinically accepted as markers of systemic inflammatory response (2, 3). Recent studies have shown that the immune response increases with a reduction in tumor load (4). Here, we present the NLR and PLR of a patient with locally advanced breast cancer who underwent a simple mastectomy to reduce the tumor load. Following surgical therapy, a remarkable regression was observed in the local recurrence area of the right mastectomy site, at the same time, the patient’s neutrophil-to-lymphocyte ratio and PLR values significantly decreased.

Keywords: Locally advanced stage, breast cancer, simple mastectomy, neutrophil-to-lymphocyte ratio, platelet-to-lymphocyte ratio

CASE PRESENTATION

A 52-year-old woman was referred to our clinic for surgical treatment of locally advanced breast cancer. From the patient’s history, it was learned that one year previously, she had presented at another center with the complaint of a mass causing color change and rash bilaterally on the breasts. The patient was diagnosed with Grade 3 invasive ductal carcinoma breast cancer on the right side. Estrogen receptor (ER) and progesterone receptor (PR) tests were strongly positive, and HER2 expression was negative (luminal A tumor). She was administered eight cycles of neoadjuvant chemotherapy, and simple mastectomy was performed on the right breast followed by hormonotherapy. It is not known why the patient underwent mastectomy of only the right breast instead of a bilateral mastectomy at the other hospital. She applied to our clinic due to increased symptoms in her left breast. When the patient presented at our clinic, the mass in her left breast had grown, there was increased rash on the breast skin, and a new mass had developed on the incision scar of the right side of the mastectomy area.

In the physical examination, multiple nodular metastatic masses, the largest of which was 1 cm in diameter, were detected on the incision scar of the mastectomy site of the right breast along with fixed lymphadenopathies in the right axilla. The left breast was completely full of tumoral mass, with tumor invasions of the skin. Inflammatory appearance of the skin, satellite nodules, and fixed lymphadenopathies in the left axilla were observed. The preoperative images are shown in Figure 1.

No abnormal findings were seen in the PA pulmonary radiograph, abdominal pelvic ultrasonograph (USG), or bone scintigraph. No pathology was determined by routine biochemistry. The patient was informed, and a simple mastectomy was applied to the left breast to reduce the tumor load. As the skin was completely covered with the tumor, the skin borders close to the sternum could not be approached with the primary incision; after removal of the breast, which was infected and had tumor necrosis, sec-
Secondary healing was applied. After 10 days, as the granulation tissue was seen to be sufficient, the open wound was repaired with graft application. No postoperative complications developed, and the patient was discharged on postoperative day 7. After removal of the tumor load in the left breast, a regression in local recurrence sites in the right mastectomy area was observed. The healed left side graft area and the regression of local recurrences on the right side are shown in Figure 2. In addition, the preoperative axillary fixed lymphadenopathies were determined to have receded on physical examination and USG imaging, and the lymph nodes were observed to have replaced the thin cortex with a hyperechogenic fatty hilus (Figure 3).

In the histopathological examination of the specimen, a Grade 3 (Scarf Bloom Richardson) invasive ductal carcinoma 11 cm in diameter was determined. There were perineural and lymphovascular invasions, many satellite tumor nodules around the tumor that had led to ulceration and an appearance of infiltration to the epidermis and breast head/areola, continuation of the tumor with a deep surgical border, and infiltration to the striated muscle in this area. ER was determined as 80%, PR as 5% positive and c-erb B-2 as negative. In the postoperative period, aromatase inhibitor was administered. The results of consecutive complete blood tests showed that the NLR and PLR values of 7.05 (N: 84%, L: 11.9%), and 407.5 (P: 326, L: 0.8×10⁹/L), respectively, on preoperative Day 1 decreased to 4.35 (N: 74%, L: 17%) and 380 (P: 266, L: 0.7×10⁹/L), respectively, on postoperative Day 5 and to 2.63 (N: 65.2%, L: 24.7%) and 251 (P: 277, L: 1.1×10⁹/L), respectively, on postoperative Day 14.

Written informed consent was obtained from the patient who participated in this case.

DISCUSSION
In cancer patients, the clinical prognosis is as much related to the properties of the tumor as to the patient’s response to the tumor (5). In recent studies, cancer and immunology have become increasingly important. For many years, immunologists have stated that the immune response is suppressed in cancer patients and the response to the tumor is reduced (6). Especially in patients with excessive tumor loads, there is evident immune suppression; with reduction of the tumor load with primary surgery, the immune response has been seen to increase. This is explained by changes occurring in T-cells including increase of antitumor responses with the reduction in tumor load (4).

Neutrophil-to-lymphocyte ratio, which can be obtained from a peripheral full blood count, is a routine clinical marker of systemic inflammatory response (2, 3). Increased peripheral neutrophils before treatment and decreased lymphocyte count have been shown to have negative effects on the survival of cancer patients (7). In addition, it has been found that just as increased neutrophil numbers are related to the paraneoplastic activity of the tumor, reduced lymphocyte numbers are related to suppression of the immune system. While neutrophils, which affect tumor growth, are the main source of angiogenesis and growth factors, lymphocytes form the patient immune response with cytotoxic cell death and production of cytotoxin, which prevent proliferation of tumor cells (8). Increased NLR in breast cancer has been shown to have increased mortality (2). In addition, in a recent study, high NLR in breast cancer patients was found to be related to lymph node metastasis (3).

Although the effects of NLR on survival in breast cancer have been widely accepted as a prognostic factor, there are few studies supporting the use of PLR as an independent prognostic factor. However, experimental studies and clinical data have shown that platelet activation is a feature of cancer, with support of neoangiogenesis, destruction of the extracellular...
matrix, and expression of adhesion molecules and growth factors (9). Seeretis et al. (10) determined a relationship between increased PLR and metastatic lymph nodes in cancer patients.

In the case presented here, the tumor load was reduced with a simple mastectomy. When the blood values of the patient were examined during this procedure, a continuous reduction in NLR was recorded together with the reduction in tumor load. The regression observed in the masses in the chest wall of the patient can be related to the decrease in the NLR value. Even though no absolute relationship has yet been shown between PLR and breast cancer, the regression of the tumor in this case together with decreased PLR values suggests that PLR can be effective in tumor prognosis.

CONCLUSION
In patients with locally advanced breast cancer, decreases in NLR and PLR values parallel to the reduction of tumor load with surgery may cause an increase in immune response and regression of the tumor. Further studies with an extensive series are required to better understand this relationship.

Informed Consent: Written informed consent was obtained from patient who participated in this study.

Peer-review: Externally peer-reviewed.


Conflict of Interest: The authors have no conflicts of interest to declare.

Financial Disclosure: The authors declared that this study has received no financial support.

REFERENCES