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CASE REPORT

# High-grade glioma in a patient with breast cancer



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#### **KEYWORDS**

anaplastic oligodendroglioma; breast cancer; high-grade glioma; metastatic brain lesions Summary Breast cancer is one of the most common origins of metastatic lesions in the central nervous system. Many patients with a breast cancer and concurrent brain tumor(s) were diagnosed to have a metastatic lesion or lesions in the brain, based exclusively on their image findings without further pathologic verification, and received radiotherapy alone thereafter. It is, however, possible that a different pathology such as primary brain malignancy, which actually warrants a specific treatment modality, may occur in such patients with an already known malignancy. We, herein, reported a 61-year-old female patient who suffered from an anaplastic oligodendroglioma 1 year after her diagnosis of breast cancer. Demographic data, characteristic imaging findings, treatment, and outcome of the patient were discussed. Copyright © 2012, Asian Surgical Association. Published by Elsevier Taiwan LLC. All rights reserved.

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#### 1. Introduction

Breast cancer is one of the most common origins of the metastatic lesions in the central nervous system (CNS). As estimated, approximately 10–15% of breast cancer patients suffer from the CNS metastatic lesion(s) throughout the course of treatment. Many patients with a breast cancer

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and concurrent brain tumor(s), however, do not receive a surgical extirpation or biopsy for the CNS lesions, because characteristic radiological findings such as a ring-like enhancing lesion or multiple lesions with peritumoral edema in the computed tomography (CT) and magnetic resonance imaging (MRI) often have made a putative diagnosis of malignant metastatic lesion or lesions. These patients are often treated with whole-brain radiotherapy (WBRT) to the brain plus systemic chemotherapy for local tumoral control, because most patients with metastatic lesions, especially those with multiple small lesions (less than  $2 \times 2 \times 2$  cm<sup>3</sup>) that originated from breast cancer, response well to the radiation treatment. 1,2 It is, however, possible that primary brain malignant gliomas, although relatively rare in incidence, may develop in patients with a systemic malignancy, and such patients actually need a treatment modality already developed and tailored for primary CNS malignancy.<sup>3</sup> We reported a female patient who suffered from an anaplastic oligodendroglioma 1 year apart after her diagnosis of breast cancer. Demographic data, characteristic imaging findings, tailored treatment, and final outcome of the patient were discussed.

#### 2. Case report

A 61-year-old woman was diagnosed to have a right-sided invasive ductal carcinoma ( $T_2N_2M_0$ , stage IIIA) and left-sided ductal carcinoma *in situ* 1 year prior to her neurologic evaluation. She had no family history of breast cancer, and had undergone right-side modified radical mastectomy and left-side total mastectomy thereafter. The tumor was confirmed to be an adenocarcinoma that was strongly positive for estrogen and progesterone receptors but was negatively stained for human epidermal growth factor receptor-2 (HER2). The patient was further treated with local radiotherapy (50 Gy for right chest wall and 50 Gy for right supraclavicular lymph nodes) and chemotherapy with

cyclophosphamide, doxorubicin, and paclitaxel, without notable adverse effects.

The patient experienced progressive recent and remote memory impairment and cognitive dysfunction 1 year after her diagnosis of breast cancer. She was also found to have mild weakness (motor grade = 4+/5) of the right upper and lower limbs. Cranial functions and other neurologic examinations were unremarkable. Further CT and MRI examinations of the brain demonstrated that the patient had multiple brain lesions, including a large  $(4 \times 4 \times 5 \text{ cm}^3 \text{ in})$ size) brain tumor located at left parieto-occipital subcortical regions extending into the ipsilateral splenium of the corpus callosum, and two other small, but apart, lesions in the lateral ventricle wall and the parietal cortex of the ipsilateral brain (Fig. 1). The patient was diagnosed to have a multifoci primary brain malignancy rather than secondary metastases, because the largest brain lesion was relatively infiltrative and deeply invasive to the corpus callosum. She subsequently underwent a craniotomy for gross total resection of the largest brain lesion. Pathologic examination demonstrated the tumor to be an anaplastic oligodendroglioma with plenty of cells having hyperchromatic, pleomorphic nuclei and abundant pale eosinophilic cytoplasm, and with vascular proliferation (Fig. 2A). Further immunohistochemical examinations showed that the tumor was strongly positive for glial fibrillary acidic protein (GFAP) but was negative for cytokeratin staining (Fig. 2B and 2C). Moreover, the Ki-67 labeling index was found to be approximately 40% (Fig. 2D), indicating a relatively high mitotic activity. Furthermore, the methylation-sensitive polymerase chain reaction (PCR) of the tumoral tissues demonstrated the O<sub>6</sub>-methylguanine-DNA methyltransferase (MGMT) promoter of the tumor to be a methylated one (Fig. 3A). Capillary electropherograms of PCR products resulting from amplification of microsatellite loci on chromosomes 1p and 19g revealed that the tumor samples contained 1p/19q codeletion (Fig. 3B). The patient then received external beam radiotherapy up to 60 Gy with concurrent daily temozolomide treatment (75 mg/kg),

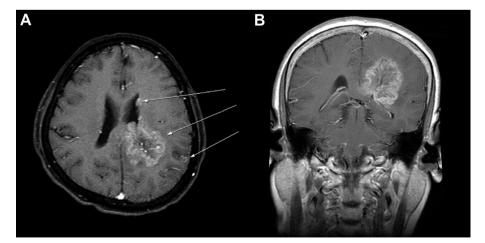


Figure 1 Magnetic resonance imaging of the patient's brain. (A) Axial T1-weighted image with gadolinium enhancement shows a large left-sided parieto-occipital lesion, located at the subcortical region but extending into the ipsilateral corpus callosum, and two concurrent small lesions in the ipsilateral lateral ventricle wall and parietal cortex (arrows). (B) Coronal section of the largest lesion shows compression of the left lateral ventricle and extension into the corpus callosum.

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