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Case Report

A rare case of breast carcinoma metastasis into a meningioma in a 64-year-old female patient^{☆,☆☆}

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ABSTRACT

This report discusses the occurrence of tumor-to-tumor metastasis—an atypical phenomenon in oncology where a secondary malignancy develops within an existing primary tumor. The case of a 64-year-old woman is presented, who, with a history of stage II invasive ductal carcinoma of the breast treated with mastectomy and chemoradiotherapy, developed neurological symptoms indicative of a secondary brain tumor. MRI and subsequent histopathological analysis post-craniotomy confirmed a meningioma with a metastatic breast carcinoma, demonstrating the clinical importance of considering tumor-to-tumor metastasis in similar patient histories.

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Introduction

Tumor-to-tumor metastasis represents a unique and complex clinical scenario characterized by a metastatic tumor emerging within another primary neoplasm [1]. Among recipient tumors, meningiomas are the most common, with breast carcinoma being a frequent metastasizing primary tumor. Al-

though exceedingly rare, this phenomenon is documented sparingly within the medical literature.

Case presentation

A 64-year-old female with a background of stage II Left breast invasive ductal carcinoma that was Estrogen and

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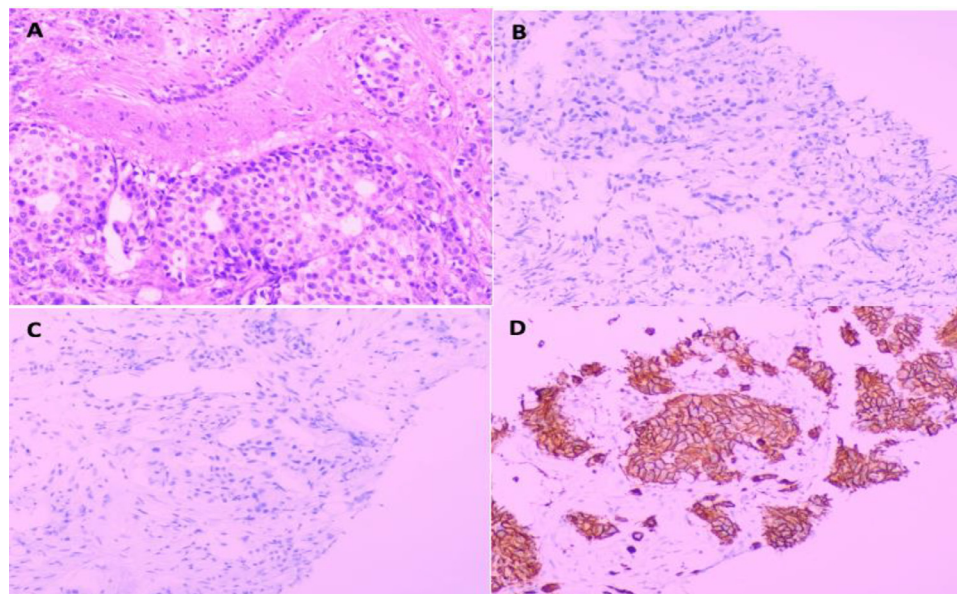


Fig. 1 – Histopathology of left breast intraductal carcinoma (IDC). (x^A) Primary carcinoma of the breast, Grade 2 (x100*) HE (y^B) estrogen receptor negative(x100*). (y^C) Progesterone Receptor negative (x100*) (y^D) Strong HER2 membrane staining (3+) ID (x100*). x-H&E Staining, y-Immunohistochemistry, * -magnification.

Progesterone receptors negative and HER-2 receptor positive, had undergone left modified radical mastectomy and adjuvant chemoradiotherapy with a good clinical response (Fig. 1). After 2 years, she developed new onset headaches and two episodes of generalized tonic-clonic seizures. Neurological examination revealed subtle Rt sided weakness with hyperreflexia and no other deficits. An MRI of the brain demonstrated a left temporal dural-based contrast-enhancing brain tumor, indicative of meningioma (Fig. 2).

The histopathology, following a gross total resection of the tumor via craniotomy, revealed two adjacent tumor types without a distinct demarcation. On immunohistochemistry, the predominant tumor was GATA3 positive, aligning with a breast primary, and exhibited the same biomarker status as the initial breast tumor. The secondary tumor displayed characteristics consistent with a meningioma.

On immunohistochemistry, the predominant tumor was GATA3 positive, in keeping with a breast primary. It also showed ER/PR negative and HER2 positive (3+) biomarker status, as was the case with the original breast tumor. The second tumor was positive for PR and S100, in a pattern consistent with a meningioma, World Health Organization (WHO) grade I meningioma (Fig. 3).

The patient's recovery was uneventful, and she was discharged on antiepileptic medications. She underwent adjuvant whole-brain radiotherapy as part of her multidisciplinary team follow-up and remains recurrence-free 22 months postcraniotomy as shown in Fig. 4.

Discussion

Tumor-to-tumor metastasis, while exceptional, has substantial implications for disease management strategies. In this

particular case, the detection of metastasis in the meningioma potentially escalated disease staging and influenced the surgical and adjuvant treatment plan [1,2].

In our case, for instance, the disease staging could have been escalated to stage IV if metastasis was detected at the initial diagnosis. Additionally, meningiomas that harbor metastasis and extend to juxtaposed tissues may necessitate more resection margins, with adjuvant radiotherapy, as evidenced in this case [3].

Further treatment relies on the extent of resection, tumor biology and clinical variables such as location [4,5]. Meningiomas are outside the blood-brain barrier, hence may be more susceptible to chemotherapy even in partially resected or inoperable meningioma with metastasis differing from that of brain parenchymal metastasis. This noteworthy characteristic underlines the importance of personalized treatment approaches based on the metastatic patterns [2,3].

Epidemiologically the most implicated primary neoplasm in tumor-to-tumor metastasis is breast carcinoma followed by lung and squamous cell carcinomas. Approximately one fifth of solid tumors metastasize to the central nervous system with a minuscule percentage metastasizing to leptomeningeal meningiomas. Literature reveals about seventy documented cases of metastasis to meningiomas [1–3].

Histopathological examination often plays a pivotal role in disease management. In our case, the unclear demarcation between the metastasis and the meningeal parenchyma made recognition challenging, underscoring the importance of careful histopathological assessment for appropriate therapeutic planning [3].

The molecular mechanisms driving metastatic anchoring and spread along the meninges remain largely elusive. While no meningioma subtypes have increased risk of tumor-to-tumor metastasis [1,6], specific features of metastasis may facilitate the process. A common putative feature could be

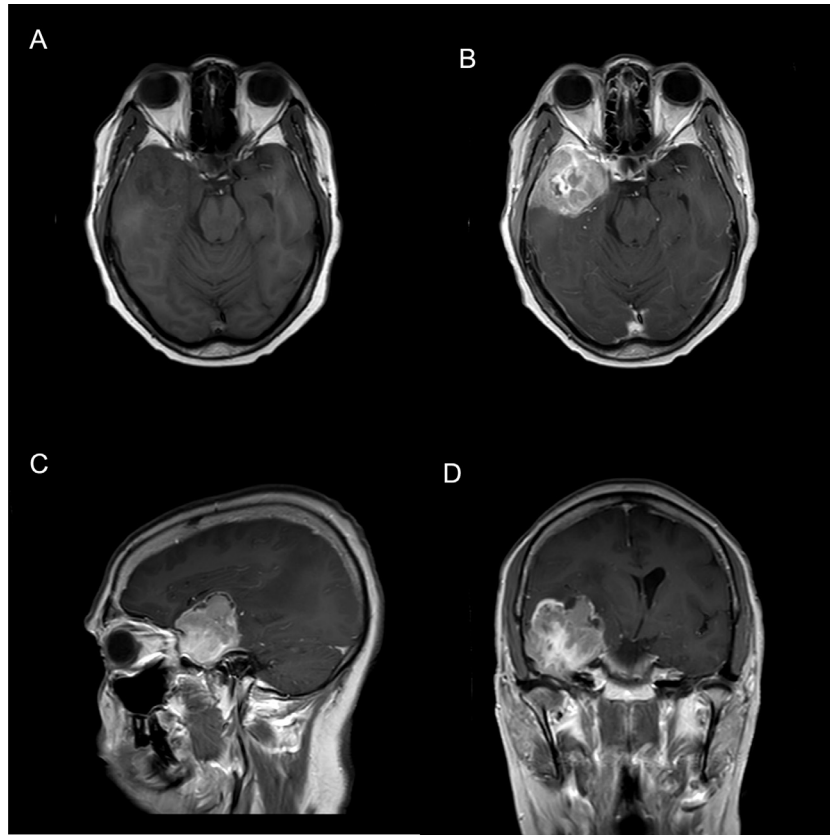


Fig. 2 – Preoperative MRI images with gadolinium. (A) MRI: Pre-Operative Axial T1 SE Non-Contrasted (B) MRI: Pre-Operative Axial T1 SE + Gadolinium Contrast (C) MRI: Pre-Operative Sagittal T1 SE + Gadolinium Contrast (D) MRI: Pre-Operative Coronal T1 SE + Gadolinium Contrast.

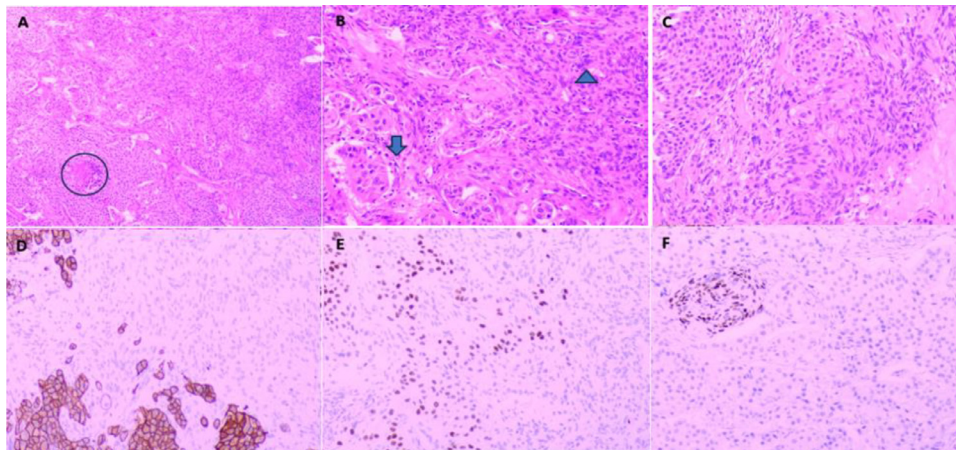


Fig. 3 – Histopathology left temporal brain tumor. (A) Comedonecrosis of the ductal carcinoma (circled) (x40*). (B) Meningotheelial cells (arrowhead), IDC cells bottom left (arrow) (x100*). (C) Meningotheelial cells, with few ductal carcinoma cells in between (circled) (x100*). (D) Strong HER2 membrane staining (3+) in IDC; negative in meningotheelial cells (x100*). (E) GATA3 nuclear positivity (brown nuclear staining) in IDC component (x100*). (F) PR positivity in meningotheelial cells, (brown nuclear staining), whorling pattern. Surrounding IDC negative (x100*). x-H&E Staining, y-Immunohistochemistry, * -magnification; IDC, intraductal carcinoma.

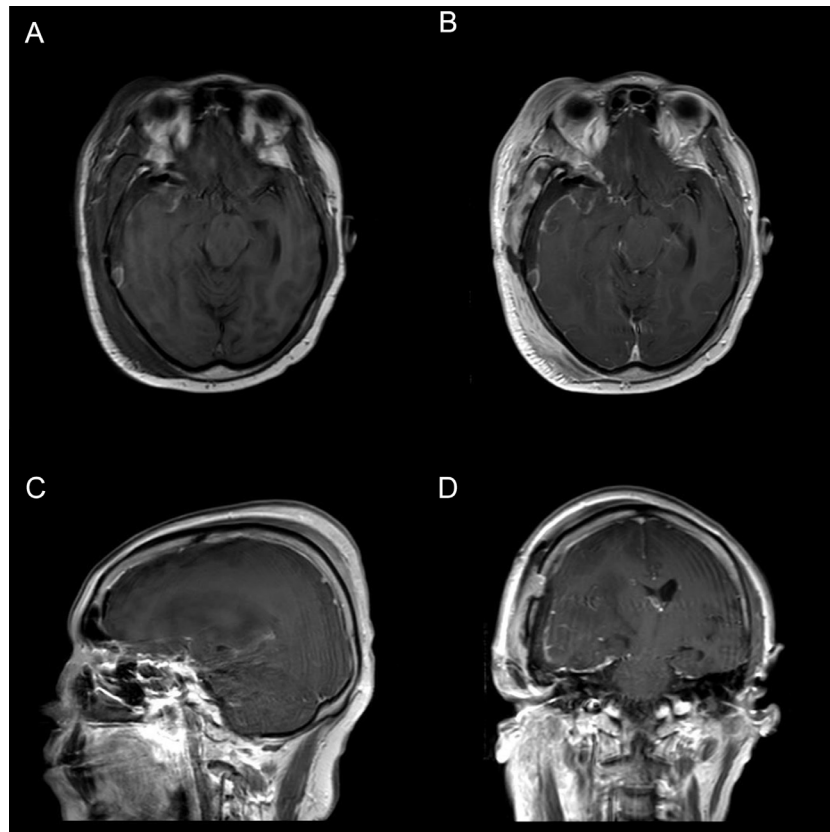


Fig. 4 – Postoperative MRI images with gadolinium. (A) MRI: Post-Operative Axial T1 SE Non-Contrasted (B) MRI: Post-Operative Axial T1 SE + Gadolinium Contrast. (C) MRI: Post-Operative Sagittal T1 SE + Gadolinium Contrast (D) MRI: Post-Operative Coronal T1 SE + Gadolinium Contrast.

hormone receptor expression and in particular progesterone receptors that seems to play a significant role in meningiomas harboring metastatic breast carcinoma. Notably our patient had a progesterone receptor-negative primary intraductal carcinoma of the breast with a progesterone receptor-positive meningioma [7,8].

Meningiomas exhibit high vascularity due to secretion of vascular endothelial growth factor (VEGF) that enhance vascular proliferation and vasogenic edema both thought to facilitate metastasis [1,9]. Multiple cell adhesion molecules like E-cadherin, P-selectin, CXCL12, ICAM, PECAM-1 and SDF-1 are implicated in aiding breast carcinoma metastases [10–13]. Additionally, the expression of mesothelin by leptomeninges and by extension meningiomas, which in itself acts as an anchoring protein that binds mucin-16 [14,15] seems to facilitate metastasis. Transmembrane mucins such as MUC1 and MUC16 ubiquitous in carcinomas are thought to facilitate metastasis of numerous carcinomas including lung adenocarcinomas [14,15]

Tumor-to-tumor metastasis have significant implications to therapeutic approaches. Clinicians must be vigilant to its possibility in patients with a known history of malignancy presenting with new neurological symptoms and have radiological evidence of meningiomas. Molecular analytical tools that are gaining traction in oncology, aiming for tailored per-

sonalized therapy depending on patient-specific factors such as genetic and molecular characteristics are likely to be the next frontier in treating these cases. Therefore, understanding the molecular basis of tumor-to-tumor metastasis may open avenues to effective personalized treatments.

A multidisciplinary team with neurosurgeons, oncologists and pathologists is mandatory for timely diagnosis and accurate treatment planning. This collaboration helps elucidate appropriate histopathological analyses with the appropriate management plan aligning with the biological features of the neoplasms.

Lastly, we must continue to broaden our insights on the mechanisms governing tumor-to-tumor metastasis. These insights will illuminate the most effective therapeutic interventions ultimately with better patient outcomes.

Conclusion

Addressing tumor-to-tumor metastasis is crucial for optimal management of cancer patients. Clinicians should maintain a high degree of vigilance for this rare occurrence, and a multidisciplinary approach is paramount in managing these complex presentations.

Patient consent

We would like to state that we have obtained written informed consent from the patient before submitting the manuscript entitled “A Rare Case of Breast Carcinoma Metastasis into a Meningioma in a 64-Year-Old Female Patient” for consideration of publication in Radiology Case Reports.

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