A 'never miss' diagnosis: Ptosis secondary to metastatic breast cancer diagnosed as involutional ptosis and a review of the literature

SAGE Open Medical Case Reports
Volume 9: 1–7
© The Author(s) 2021
Article reuse guidelines
sagepub.com/journals-permissions
DOI: 10.1177/2050313X211040680
journals.sagepub.com/home/sco

\$SAGE

Mustafa Safi¹, Katayun Fethat² and Rona Z Silkiss^{1,3}

Abstract

We present a case of ptosis secondary to metastatic lobular carcinoma of the breast which was initially diagnosed as involutional ptosis. A 67-year-old woman previously diagnosed with lobular carcinoma of the breast presented to our clinic with mild restriction of lateral gaze and persistent droopiness of her right eyelid (associated with decreased levator function) despite recent repair of a suspected involutional ptosis. Orbital magnetic resonance imaging revealed a mass in the right orbit which was biopsied and diagnosed as lobular carcinoma of the breast. Poor levator function is rarely present in involutional ptosis. Especially in conjunction with abnormal extraocular motility, other etiologies of acquired ptosis must be considered. This case highlights the importance of patient history and ocular examination in identifying the underlying etiology of ptosis. A review of the literature to evaluate the incidence of signs and symptoms associated with metastatic breast cancer to the orbit is included.

Keywords

ptosis, lobular, carcinoma, metastasis, orbit, breast

Date received: 9 April 2021; accepted: 20 July 2021

Introduction

Breast cancer is a common source of orbital metastasis in women. Orbital metastasis can present with ptosis, proptosis, periorbital edema, diplopia, among other signs.² Motility deficits can also be present when metastatic lesions involve the extraocular muscles, manifesting as 'bumpy muscles' on imaging. At times, the periocular presentation associated with an orbital metastasis may be subtle. Furthermore, the associated clinical manifestations can evolve with time; what manifests for one clinician at a given point in time may be different for the next. Herein, we present the case of a patient who presented to our clinic with persistent ptosis despite prior surgical repair for involutional ptosis. Given the history of breast cancer, ptosis manifesting with poor levator function, and restrictive lateral gaze, orbital imaging was undertaken and a neoplastic etiology was identified. With atypical cases, the patient history, clinical signs, and judicious use of orbital imaging may guide the clinician to the most likely diagnosis.² In addition to our case, a review of the literature evaluating the incidence of the signs and

symptoms, location, and histologic features of orbital breast cancer metastasis is provided.

Case report

A 67-year-old woman diagnosed with breast cancer in 2017 was referred to our clinic for persistent right upper eyelid ptosis. She was previously treated by an ophthalmologist, who diagnosed an involutional ptosis. Upon presentation, her visual acuity (VA) was 20/200 in the right eye (OD), 20/40 with eyelid elevation, and 20/40 in the left eye (OS). There was a slight decrease in abduction OD. The margin reflex distance 1 (MRD 1) was -2 mm OD and 2 mm OS

¹California Pacific Medical Center, San Francisco, CA, USA ²University of California, Davis, Davis, CA, USA ³Silkiss Eye Surgery, San Francisco, CA, USA

Corresponding Author:

Rona Z Silkiss, Silkiss Eye Surgery, FACS, 711 Van Ness Avenue Suite 340, San Francisco, CA 94102, USA.
Email: DrSilkiss@silkisseyesurgery.com



Figure 1. External photo demonstrating margin reflex distance I (MRD I) of -2 mm in the right eye and 2 mm in the left. Note the asymmetry of the superior sulcus.

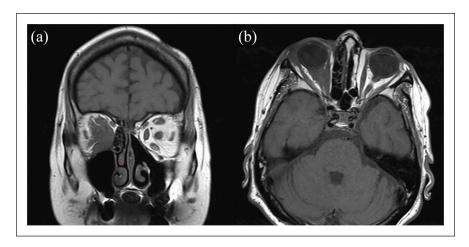


Figure 2. Coronal TI image demonstrating infiltrating soft tissue throughout the right orbit with abutment of the inferior, medial, and superior recti (a). Encasement of the optic nerve is demonstrated on both the coronal TI and axial TI images (a and b). Post-surgical changes from prior maxillary sinus surgery is present in panel a.

(Figure 1). The margin reflex distance 2 (MRD 2) was normal in both eyes (OU). Levator function was 2 mm OD and 10 mm OS. There was fullness in the right superior sulcus relative to left; the right superior sulcus was non-tender and firm upon palpation. The remainder of the eye examination was normal.

Magnetic resonance imaging (MRI) of the orbit with and without contrast demonstrated an infiltrating enhancing soft tissue mass throughout the intraconal and extraconal space with extension to the orbital apex (Figure 2). Abutment of the superior, medial, and inferior recti with associated encasement of the optic nerve was noted (Figure 2). This

likely contributed to the restriction of the right eye in lateral gaze. Within a week of presentation, a right anterior orbitotomy with biopsy via a lower lid transconjunctival approach was performed. A gray-white mass was visualized intraoperatively and biopsied. The biopsy demonstrated lobular carcinoma (Figure 3). Our patient was referred to oncology and radiation therapy for further management and treatment.

Methods

A comprehensive review of the literature was performed using PubMed and Google Scholar. The search phrase

Safi et al. 3

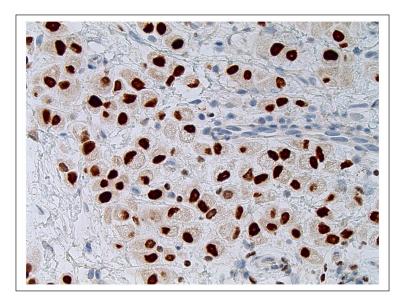


Figure 3. Hematoxylin and eosin stain ($40 \times$ magnification) with immunoperoxidase stain for GATA-3, transcription factor, expressed in breast carcinoma.

Table 1. Frequency of age, gender and laterality of orbital metastasis.

Age	n	(%)	Gender	n	(%)	Laterality ^a	n	(%)
			Female			0		50.0
50<	65	64.4	Male	2	2.0	Left	58	50.0

^a I5 cases were bilateral.

'breast cancer metastasis to the orbit', 'metastatic breast cancer to orbit', and 'orbital metastasis from breast cancer' identified 101 cases.^{3–95} Articles were researched for age, gender, and laterality of metastasis, ocular signs and symptoms, location of metastasis, and histologic features of breast cancer.

Results

A total of 101 cases and 116 eyes were identified. Increased incidence of orbital metastasis was noted in women younger than 50 years of age (n=65, 64.4%) relative to patients older than 50 (n=36, 35.6%). The right and left orbit were equally affected (Table 1). The most frequent symptoms experienced were blurred vision (37.9%), double vision (34.5%), eye pain (32.8%), bulging eye (16.4%), and droopy eyelid (10.3%). Ophthalmic examination most commonly demonstrated gaze deficits (50.9%), decreased VA (48.3%), exophthalmos (28.5%), ptosis (20.7%), and enophthalmos (14.7%). Orbital soft tissue was a frequent site of metastasis (47.4%), followed by extraocular muscle (43.1%), optic nerve (30.2%), orbital bone (20.1%), and lacrimal gland (3.5%). Invasive lobular carcinoma (33.7%) had the highest incidence of orbital metastasis followed by invasive ductal carcinoma (28.7%) (Table 2).

Discussion

Routine involutional ptosis presents with normal levator function. The presence of decreased levator function as well as double vision or other newly acquired neurologic signs should alert the clinician that the patient may not have a routine involutional ptosis. A sinister and potentially life-threatening condition such as an intraorbital or intracranial tumor may exist. Additionally, the patient should be queried regarding a history of thyroid eye disease, diabetes, and myasthenia gravis. These conditions can also lead to ptosis with decreased levator function.

Consistent with our review, our patient presented with complaints of a persistent droopy eyelid and was found to have abnormal extraocular movements and ptosis on examination, symptoms and signs which can frequently be seen in association with metastatic breast cancer to the orbit. Metastasis to the orbital soft tissue, extraocular muscle, and optic nerve are the most common sites of orbital metastasis from breast cancer. This is likely due to the rich blood supply these structures possess which allows for hematogenous seeding. Our patient demonstrated involvement of all three structures.

Evaluation of the orbit is best achieved with MRI which allows for enhanced visualization of soft tissues without the radiation exposure of computed tomography (CT) which provides greater bony rather than soft tissue details. The MRI of this patient demonstrated the presence of an intraorbital mass. This was biopsied and diagnosed as invasive lobular carcinoma of the breast. Orbital metastasis occurs via the hematogenous route. The finding of orbital metastasis is suggestive of malignant seeding to other organs as well. A multidisciplinary team, including an oncologist, radiation oncologist, pain specialist, ophthalmologist, and ancillary

7.9

1.0

1.0

1.0

1.0

Symptoms	nª	(%)	Signs	nª	(%)	Location of orbital metastases	nª	(%)	Histology	n	(%)
Blurred vision	44	37.9	Gaze deficits	59	50.9	Soft tissue	55	47.4	Invasive lobular	34	33.7
Double vision	40	34.5	Decreased visual acuity	56	48.3	Extraocular muscle	50	43.I	Invasive ductal carcinoma	29	28.7
Pain	38	32.8	Exophthalmos	33	28.5	Optic nerve	35	30.2	Unknown ^b	26	25.7

24 20.7 Bone

15 12.9

13 11.2

12 10.3

12 10.3

17 14.7 Lacrimal gland

Table 2. Frequency of signs, symptoms, location of orbital metastases, and histology of breast cancer.

1.7 Optic disc swelling

10.3 Enophthalmos

4.3 Palpable mass

3.6 Iris/pupillary abnormalities

16.4 Ptosis

10.3 Swelling

Bulging eye

Swelling

Redness

Dryness

Droopy eyelid

Eyelid thickening

19

12

12

5

3

services, should take a holistic approach focusing on prolonging survival and the palliation of symptoms. Hormone therapy, chemotherapy, immunotherapy, and radiation treatment are options available in the management of patients with orbital metastasis.⁸⁸

Conclusion

We present a case of acquired ptosis with poor levator function and abduction deficit associated with metastatic lobular carcinoma of the orbit. In atypical ptosis cases, a thorough history and physical examination with close follow-up is needed. MRI is the imaging modality of choice when an intraorbital and/or intracranial neoplastic process is suspected. Management requires a multidisciplinary team and approach.

Acknowledgements

We would like to extend our gratitude to the Highland Hospital medical librarian, Laurie Bagley, who was of great assistance to our study.

Declaration of conflicting interests

The author(s) declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

Ethical approval

Our institution does not require ethical approval for reporting individual cases or case series.

Funding

The author(s) disclosed receipt of the following financial support for the research, authorship, and/or publication of this article: Funding was provided by the California Pacific Medical Center, San Francisco, CA.

Informed consent

Written informed consent was obtained from the patient(s) for their anonymized information to be published in this article.

ORCID iD

Mustafa Safi https://orcid.org/0000-0002-6773-0038

References

 Shields JA, Shields CL and Scartozzi R. Survey of 1264 patients with orbital tumors and simulating lesions: the 2002 Montgomery lecture, part 1. *Ophthalmology* 2004; 111(5): 997–1008.

24 20.1 Scirrhous carcinoma

3.5 Eccrine adenocarcinoma

Phyllodes tumor

Inconclusive

Mucinous carcinoma

- 2. Allen RC. Orbital metastases: when to suspect? When to biopsy? *Middle East Afr J Ophthalmol* 2018; 25(2): 60–64.
- Aghdam KA, Zand A and Sanjari MS. Isolated unilateral infiltrative optic neuropathy in a patient with breast cancer. *Turk J Ophthalmol* 2019; 49(3): 171–174.
- 4. Ahmad W, Ahmad U and Hassan A. Retrobulbar metastasis of breast primary: once in a blue moon lesion. *J Pak Med Assoc* 2020; 70(12(A)): 2294–2295.
- Allaire GS, Corriveau C and Arbour JD. Metastasis to the optic nerve: clinicopathological correlations. *Can J Ophthalmol* 1995; 30(6): 306–311.
- Archilei G. Metastasi orbito-parananasale da carcinoma mammario [Orbito-paranasal metastases of breast carcinoma]. *Boll Mal Orecch Gola Naso* 1959; 77(3): 218–229.
- Asproudis I, Gorezis S, Charalabopoulos K, et al. Breast carcinoma metastasis to the orbit and paranasal sinuses: a case report. Exp Oncol 2004; 26(3): 246–248.
- 8. Atasoy BM, Cetin IA, Bozkurt SU, et al. Metastasis to paranasal sinuses and orbita of breast cancer with a rare metachronous tumor of the uterine cervix. *J Craniofac Surg* 2013; 24(1): e64–e65.
- Backhouse O, Simmons I, Frank A, et al. Optic nerve breast metastasis mimicking meningioma. Aust N Z J Ophthalmol 1998; 26(3): 247–249.
- 10. Bagnolesi P, Laddaga F, Mancino M, et al. Metastasi orbitarie da carcinoma mammario. Quadro radiologico [Orbital metastasis of breast carcinoma: radiologic picture]. *Radiol Med* 1994; 88(4): 483–486.
- 11. Cherekaev VA, Lasunin NV, Stepanian MA, et al. Breast carcinoma metastasis to the optic nerve: case report and review of literature. *Zh Vopr Neirokhir Im N N Burdenko* 2013; 77(3): 42–48; discussion 48.
- 12. Chew C, Wan Hitam WH and Ahmad Tajudin LS. Leptomeningeal carcinomatosis with optic nerve metastasis secondary to breast cancer. *Cureus* 2021; 13(3): e14200.

^aEach case can have more than one sign, symptom, or area of orbital involvement.

^bHistology not available.

Safi et al. 5

 Cho HK, Park SH and Shin SY. Isolated optic nerve metastasis of breast cancer initially mimicking retrobulbar optic neuritis. *Eur J Ophthalmol* 2011; 21(4): 513–515.

- Coutinho I, Marques M, Almeida R, et al. Extraocular muscles involvement as the initial presentation in metastatic breast cancer. *J Breast Cancer* 2018; 21(3): 339–342.
- Danek DJ, Blessing NW and Tse DT. Corneal perforation with uveal prolapse: an initial presentation of orbital metastatic breast cancer. Am J Ophthalmol Case Rep 2019; 16: 100551.
- Dieing A, Schulz CO, Schmid P, et al. Orbital metastases in breast cancer: report of two cases and review of the literature. *J Cancer Res Clin Oncol* 2004; 130(12): 745–748.
- Eckardt AM, Rana M, Essig H, et al. Orbital metastases as first sign of metastatic spread in breast cancer: case report and review of the literature. *Head Neck Oncol* 2011; 3: 37.
- El Bakraoui K and El Morabit B. Orbital metastasis from triple-negative breast cancer: case report and literature review. *Case Rep Oncol* 2020; 13(2): 1042–1046.
- El-Khazen Dupuis J, Marchand M, Javidi S, et al. Enophthalmos as the initial systemic finding of undiagnosed metastatic breast carcinoma. *Int Med Case Rep J* 2021; 14: 25–31.
- Framarino-Dei-Malatesta M, Chiarito A, Bianciardi F, et al. Metastases to extraocular muscles from breast cancer: case report and up-to-date review of the literature. *BMC Cancer* 2019; 19(1): 36.
- Furuno K, Asaga T, Uchiyama M, et al. Orbital and stomach metastasis from invasive lobular breast carcinoma. *Nippon Ganka Gakkai Zasshi* 1998; 102(2): 135–141.
- Fyrmpas G, Televantou D, Papageorgiou V, et al. Unsuspected breast carcinoma presenting as orbital complication of rhinosinusitis. *Eur Arch Otorhinolaryngol* 2008; 265(8): 979–982
- Gaddi MJS, Yuga ACQ, Dofitas RB, et al. Surgery for orbital metastasis from breast carcinoma initially presenting with progressive proptosis. *BMJ Case Rep* 2020; 13(12): e237158.
- Gasperini J, Black E and Van Stavern G. Perineural metastasis
 of breast cancer treated with optic nerve sheath fenestration.

 Ophthalmic Plast Reconstr Surg 2007; 23(4): 331–333.
- Giffon D, Ravault P and Trepsat C. Métastase orbito-palpébrale d'un cancer du sein [Orbitopalpebral metastasis of breast cancer]. *Bull Soc Ophtalmol Fr* 1987; 87(5): 701–702.
- Glazer LC, Harris GJ and Simons KB. Orbital metastasis as the presenting sign of adenocarcinoma of the breast. *Ophthalmic Plast Reconstr Surg* 1991; 7(4): 252–255.
- Gonçalves AC, Moura FC and Monteiro ML. Bilateral progressive enophthalmos as the presenting sign of metastatic breast carcinoma. *Ophthalmic Plast Reconstr Surg* 2005; 21(4): 311–313.
- González F and López-Couto C. Metástasis orbitarias. Serie de cuatro casos y revisión de la literatura [Orbital metastases: a report of four cases and a review of the literature]. Arch Soc Esp Oftalmol 2006; 81(8): 451–462.
- Gupta S, Bhatt VR and Varma S. Unilateral orbital pain and eyelid swelling in a 46-year-old woman: orbital metastasis of occult invasive lobular carcinoma of breast masquerading orbital pseudotumour. *BMJ Case Rep* 2011; 2011: bcr1220103580.

- Harris AL, Montgomery A, Reyes RR, et al. Carcinoid tumour presenting as an orbital metastasis. *Clin Oncol* 1981; 7(4): 365–372.
- Hashimoto M, Tomura N and Watarai J. Retrobulbar orbital metastasis mimicking meningioma. *Radiat Med* 1995; 13(2): 77–79.
- Hemmanouil I, Stöver T and Schwab B. Der interessante Fall Nr. 28. Orbitale Metastasierung des bekannten Mammakarzinoms [Interesting case no. 28: orbital metastasis of established breast cancer]. *Laryngorhinootologie* 1999; 78(9): 525–526.
- Hernández Pardines F, Serra Verdú MC, Bernal Vidal A, et al. Lateral rectus muscle biopsy as diagnosis of unknown metastatic breast cancer: Biopsia del músculo recto lateral orbitario como diagnóstico de cáncer de mama metastásico no conocido. Arch Soc Esp Oftalmol 2019; 94(4): 192–195.
- 34. Hoffmann K, Löblich HJ and Weinrich W. Enophthalmus mit Motillitätsstörung als seltenes klinisches Symptom metastasierter Mammakarzinome [Enophthalmos with limitation of bulbar motility as a rare clinical symptom of metastasizing carcinoma of the breast (author's transl)]. Klin Monbl Augenheilkd 1980; 177(3): 376–379.
- 35. Huda N and Venable HP. Metastasis of carcinoma of the breast to both orbits. *Am J Ophthalmol* 1967; 64(4): 779–780.
- 36. Jacobs M and Benger R. Metastatic breast carcinoma of the orbit. *Aust N Z J Ophthalmol* 1989; 17(4): 357–361.
- Janicijevic-Petrovic M, Sarenac T, Sreckovic S, et al. Orbital metastases from breast cancer: a case report. Bosn J Basic Med Sci 2011; 11(4): 253–255.
- 38. Kadivar M, Joulaee A, Kashkouli MB, et al. Orbital metastasis as the first presentation of nonpalpable invasive lobular carcinoma of the breast. *Breast J* 2006; 12(1): 75–76.
- Kanoh T, Nakano Y, Inatome J, et al. A case of successfully treated orbital metastasis from breast cancer by radiation therapy. Gan To Kagaku Ryoho 2008; 35(12): 2231–2233.
- Kim CU and Pearce WA. Metastatic breast carcinoma involving the optic disc. Am J Ophthalmol Case Rep 2020; 18: 100698
- Kim HJ, Wojno TH and Grossniklaus H. Atypical bilateral orbital metastases of lobular breast carcinoma. *Ophthalmic Plast Reconstr Surg* 2012; 28(6): e142–143.
- 42. Kluska A, Papis-Ubych A, Fijuth J, et al. Intraorbital extraocular metastasis of breast cancer 11 years after mastectomy: case report and review of the literature. *J Obstet Gynaecol* 2019; 39(1): 126–128.
- Kokufu I, Tanei T, Taniguchi H, et al. Two cases of effective weekly paclitaxel administration and concurrent radiation for metastatic breast cancer. *Gan To Kagaku Ryoho* 2003; 30(1): 111–114.
- Kouvaris JR, Gkongkou PV, Papadimitriou CA, et al. Bilateral metastases to extraocular muscles from lobular breast carcinoma. *Onkologie* 2008; 31(7): 387–389.
- 45. Kuchel JM and Bowling JC. Bilateral lower eyelid masses. *Arch Dermatol* 2006; 142(10): 1351–1356.
- Kuo SC, Hsiao SC, Chiou CC, et al. Metastatic carcinoma of the breast: a case with the unusual presentation of unilateral periorbital edema. *Jpn J Ophthalmol* 2008; 52(4): 305–307.
- Lagrèze WD, Wesendahl TA and Kommerell G. Enophthalmus durch Orbitametastase eines Mamma-Carcinoms [Enophthalmos

- caused by orbital metastasis of breast carcinoma]. Klin Monbl Augenheilkd 1997; 211(1): 68–69.
- 48. Lin IH, Kuo BI and Liu FY. Adjuvant intravitreal bevacizumab injection for choroidal and orbital metastases of refractory invasive ductal carcinoma of the breast. *Medicina* 2021; 57(5): 404.
- 49. Lucereau-Barbier M, El Falah S, Desoutter M, et al. Un adénocarcinome mammaire découvert par une métastase orbitaire: à propos d'un cas et revue de la littérature [Breast adenocarcinoma discovered by orbital metastases: a case report and review of the literature]. *J Gynecol Obstet Biol Reprod (Paris)* 2012; 41(1): 96–99.
- Martínez-Prieto M, Sánchez Basurto C, Sánchez Forgach E, et al. Cáncer de mama con metástasis hacia las orbitas. Reporte de un caso [Breast cancer with metastases to orbits: case report]. Ginecol Obstet Mex 2007; 75(9): 553–555.
- 51. Martorell-Calatayud A, Requena C, Díaz-Recuero JL, et al. Mask-like metastasis: report of 2 cases of 4 eyelid metastases and review of the literature. *Am J Dermatopathol* 2010; 32(1): 9–14.
- Milman T, Pliner L and Langer PD. Breast carcinoma metastatic to the orbit: an unusually late presentation. *Ophthalmic Plast Reconstr Surg* 2008; 24(6): 480–482.
- 53. Mohadjer Y and Holds JB. Orbital metastasis as the initial finding of breast carcinoma: a ten-year survival. *Ophthalmic Plast Reconstr Surg* 2005; 21(1): 65–66.
- 54. Mora-Guzmán I, Maqueda González R, Doblado Cardellach B, et al. Isolated orbital metastasis as an initial presentation of a breast cancer: metástasis orbitaria solitaria como presentación inicial de cáncer de mama. *Cir Esp* 2018; 96(2): 119.
- 55. Mourits MP, Saeed P and Kloos RJ. Enophthalmus als eerste uiting van borstkanker [Enophthalmos as a first sign of breast cancer]. *Ned Tijdschr Geneeskd* 2015; 159: A9114.
- 56. Muhd H, Zuhaimy H, Ismail MF, et al. Orbital metastasis as the initial presentation of breast cancer. *Malays Fam Physician* 2020; 15(3): 74–78.
- 57. Murthy R, Gupta A, Hegde S, et al. Bilateral multiple extraocular muscle metastasis from breast carcinoma. *Indian J Ophthalmol* 2011; 59(5): 381–382.
- Nifosí G and Zuccarello M. Unilateral localized extraocular muscle metastasis by lobular breast carcinoma. *BMJ Case Rep* 2018; 2018: bcr2018224726.
- Nirmala S, Krishnaswamy M, Janaki MG, et al. Unilateral solitary choroid metastasis from breast cancer: rewarding results of external radiotherapy. *J Cancer Res Ther* 2008; 4(4): 206–208.
- 60. Oprean CM, Badau LM, Segarceanu NA, et al. Unilateral orbital metastasis as the unique symptom in the onset of breast cancer in a postmenopausal woman: case report and review of the literature. *Diagnostics* 2021; 11(4): 725.
- Orr CK, Cochran E and Shinder R. Metastatic scirrhous breast carcinoma to orbit causing enophthalmos. *Ophthalmology* 2016; 123(7): 1529.
- Papathanassiou M, Nikita E, Theodossiadis P, et al. Orbital metastasis secondary to breast cancer mimicking thyroidassociated ophthalmopathy. *Clin Exp Optom* 2010; 93(5): 368–369.
- Patel MM, Lefebvre DR, Lee NG, et al. Gaze-evoked amaurosis from orbital breast carcinoma metastasis. *Ophthalmic Plast Reconstr Surg* 2013; 29(4): e98–e101.

- 64. Peckham EL, Giblen G, Kim AK, et al. Bilateral extraocular muscle metastasis from primary breast cancer. *Neurology* 2005; 65(1): 74.
- 65. Pinto Proença R, Fernandes J, Burnier MN, et al. Orbital metastasis from an occult breast carcinoma (T0, N1, M1). *BMJ Case Rep* 2018; 2018: bcr2017223542.
- 66. Polito E, Pichierri P, Occhini R, et al. Orbital metastasis associated with primary breast carcinoma in a man detected during peribulbar anesthesia for cataract surgery. *Eur J Ophthalmol* 2008; 18(6): 1031–1033.
- 67. Porcel JM, Salud A and Balil A. Exoftalmos unilateral secundario a metástasis orbitaria como manifestación inicial de un cáncer de mama [Unilateral exophthalmos secondary to orbital metastasis as initial manifestation of breast cancer]. *An Med Interna* 2002; 19(3): 159–160.
- 68. Radnot M and Varga M. Metastatic tumor of the orbit. *Ann Ophthalmol* 1975; 7(11): 1465–1467.
- 69. Rajabi MT, Hassanpoor N, Parsa R, et al. Spontaneous retrobulbar hemorrhage in a patient with breast cancer: a case report. *J Curr Ophthalmol* 2016; 28(1): 48–51.
- Rakul Nambiar K, Ajith PS and Arjunan A. Unilateral proptosis as the initial manifestation of malignancy. *J Egypt Natl Canc Inst* 2017; 29(3): 159–161.
- Razem B and Slimani F. An early orbital metastasis from breast cancer: a case report. *Int J Surg Case Rep* 2021; 78: 300–302.
- 72. Reeves D, Levine MR and Lash R. Nonpalpable breast carcinoma presenting as orbital infiltration: case presentation and literature review. *Ophthalmic Plast Reconstr Surg* 2002; 18(1): 84–88.
- Reyntjens I, De Groot V, Van Marck E, et al. Biopsy in single extraocular muscle hypertrophy. *Bull Soc Belge Ophtalmol* 1998; 268: 163–169.
- Rogers LR, Remer SE and Tejwani S. Durable response of breast cancer leptomeningeal metastasis to capecitabine monotherapy. *Neuro Oncol* 2004; 6(1): 63–64.
- 75. Rossi L, Zancla S, Civitelli L, et al. An unusual orbital metastasis of breast cancer. *Breast Dis* 2014; 34(4): 173–176.
- Saffra N, Rakhamimov A, Wrzolek MA, et al. Orbital metastasis as the initial presentation in bilateral lobular invasive carcinoma of the breast. *Ophthalmic Plast Reconstr Surg* 2014; 30(2): e30–12.
- 77. Saitoh A, Amemiya T and Tsuda N. Metastasis of breast carcinoma to eyelid and orbit of a postmenopausal woman: good response to tamoxifen therapy. *Ophthalmologica* 1997; 211(6): 362–366.
- 78. Sakurai K, Amano S, Enomoto K, et al. A case of advanced breast cancer with meningeal carcinomas and orbital metastasis successfully treated with multi-disciplinary therapy. *Gan To Kagaku Ryoho* 2006; 33(12): 1913–1915.
- 79. Salinas-Botrán A and Guarín-Corredor MJ. Orbital metastasis in breast cancer. *N Engl J Med* 2019; 381(12): e26.
- 80. Sánchez Orgaz M, Gonzalez Pessolani T, Pozo Kreilinger JJ, et al. Orbital and conjunctival metastasis from lobular breast carcinoma. *Orbit* 2017; 36(4): 197–200.
- 81. Solari HP, Ventura MP, Cheema DP, et al. Orbital metastasis from breast carcinoma presenting as neurotrophic keratitis. *Can J Ophthalmol* 2006; 41(1): 93–96.
- 82. Surace D, Piscioli I, Morelli L, et al. Orbital metastasis as the first sign of 'dormant' breast cancer dissemination 25 years after mastectomy. *Jpn J Ophthalmol* 2008; 52(5): 423–425.

Safi et al.

Talwar V, Vaid AK, Doval DC, et al. Isolated intraorbital metastasis in breast carcinoma. *J Assoc Physicians India* 2007; 55: 451–452.

- Tamai H, Ishida K, Murakami K, et al. Compression neuropathy caused by cancer metastasis to the optic nerve canal. *BMC Res Notes* 2013; 6: 546.
- Tiwari V, Pande SC, Verma K, et al. Paranasal sinus and retroorbital metastasis in a case of breast carcinoma: a clinicoradiological review. *BMJ Case Rep* 2014; 2014: bcr2014204094.
- Toller KK, Gigantelli JW and Spalding MJ. Bilateral orbital metastases from breast carcinoma: a case of false pseudotumor. *Ophthalmology* 1998; 105(10): 1897–1901.
- 87. Tomizawa Y, Ocque R and Ohori NP. Orbital metastasis as the initial presentation of invasive lobular carcinoma of breast. *Intern Med* 2012; 51(12): 1635–1638.
- 88. Van der Heijden A, Twijnstra A, Lamers WP, et al. An unusual cause of diplopia in a cancer patient. *Eur J Cancer* 1991; 27(10): 1315–1316.
- 89. Vempuluru VS, Mishra DK and Kaliki S. Malignant phyllodes tumor of the breast with metastasis to the orbit: a rare case report. *Ophthalmic Plast Reconstr Surg* 2021; 37(1): e5–e7.

- Vlachostergios PJ, Voutsadakis IA and Papandreou CN. Orbital metastasis of breast carcinoma. *Breast Cancer (Auckl)* 2009; 3: 91–97.
- Weiss R, Grisold W, Jellinger K, et al. Metastasis of solid tumors in extraocular muscles. *Acta Neuropathol* 1984; 65(2): 168–171.
- 92. Wolstencroft SJ, Hodder SC, Askill CF, et al. Orbital metastasis due to interval lobular carcinoma of the breast: a potential mimic of lymphoma. *Arch Ophthalmol* 1999; 117(10): 1419–1421.
- 93. Wu ZY and Mo YQ. Orbital metastasis-report of 5 cases. *Yan Ke Xue Bao* 1987; 3(3): 156–159.
- Yabaş Kızıloğlu Paksoy Türköz ÖF, Totuk Gedar ÖM, Mestanoğlu M, et al. Breast carcinoma metastasis to the medial rectus muscle: case report. *Turk J Ophthalmol* 2019; 49(3): 168–170.
- Zhang GJ, Adachi I, Yin DF, et al. Eyelid metastasis from breast cancer showing marked response to chemotherapy. *Jpn J Clin Oncol* 1995; 25(1): 10–15.
- Farber SE and Codner MA. Evaluation and management of acquired ptosis. Plast Aesthet Res 2020; 7: 20.