



DOI: 10.32768/abc.202073137-142

Large Malignant Phyllodes Tumor of the Breast: Case Report of a Young Female and Review of Literature

Melissa Kyriakos Saad^{*a}, Imad El Hajj^a, Elias Saikaly^a^a Saint George Hospital University Medical Center, Beirut, Lebanon

ARTICLE INFO

Received:

15 June 2020

Revised:

21 July 2020

Accepted:

29 July 2020

Key words:Phyllodes tumor,
breast cancer,
management

ABSTRACT

Background: Phyllodes tumors are rare fibroepithelial neoplasm of the breast, representing 0.3 to 0.5% of all female breast neoplasms. The term “phyllodes tumor” by the World Health Organization (WHO) categorizes it into benign, borderline, and malignant tumors based on histopathological characteristics. Malignant phyllodes tumor is an uncommon but aggressive breast malignancy and accounts for approximately 25% of all phyllodes tumors.

Case Presentation: A 23-year-old female patient Gravida 0 Para 0, previously healthy, with no family history of breast or ovarian cancer was referred to our institution for managing a right breast mass increasing in size over a period of 1 month associated with reddish-brown discoloration of the skin. Breast ultrasound showed an isodense oval mass measuring approximately 16x14x12cm in the right lower outer quadrant of her breast. Core biopsy and FNA done revealed a low grade phyllodes tumor. MRI of breasts showed a huge mass of the right breast occupying all quadrants, measuring 15x14.5 cm in its greatest axis with involvement of the pectoralis major muscle, with no evidence of suspicious axillary lymphadenopathy. Distant metastatic work-up with CT scan of chest abdomen and pelvis and a PET scan did not show distant metastasis. Hence, the patient underwent right total mastectomy with a final pathology compatible with malignant phyllodes tumor. She is on regular follow up and 18 months post-surgery is still disease free.

Conclusion: Management of malignant phyllodes tumor remains debatable, especially when it comes to the effect of adjuvant radiotherapy and chemotherapy. From a surgical perspective, whether phyllodes tumors should be considered as epithelial breast cancers or as soft tissue sarcoma is another debate. On the other hand, although challenging, accurate identification of phyllodes tumor initially may aid in decreased recurrence.

Introduction

Phyllodes tumors are uncommon fibroepithelial neoplasm of the breast, accounting for 0.3 to 0.5% of all female breast neoplasms,¹ with an incidence rate of about 2.1 per million mostly occurring in women aged 45 to 49 years.^{2,3} The term “phyllodes tumor” by the World Health Organization (WHO) has

categorized it into benign, borderline, and malignant tumors on the basis of histopathological characteristics. Malignant phyllodes tumor is an uncommon but aggressive breast malignancy and constitutes nearly 25% of all phyllodes tumors. Although malignant phyllodes tumor typically occurs in the fourth to sixth decades of life, it can also occur in younger women. In a study by Abusalem et al., it was estimated that one out of 26 phyllodes tumor patients was malignant in patients 20-29 years of age.^{4,5} Patients usually present with a spectrum of complaints starting with a palpable mass or abnormal mammogram and reaching architecture

Address for correspondence:

Melissa Kyriakos Saad, MD

Address: General surgery department, Saint Georges Hospital University Medical Center, Ashrafieh, Beirut, P.O.Box 166 378, Lebanon

Email: melissa.kyriakossaad@gmail.com



Figure 1. Patient with right breast malignant phyllodes (a. supine view; b. lateral view)

distortion of the breast. Accurate preoperative diagnosis by clinical, radiological and histological examination is of paramount importance as this will allow optimal surgical planning and, hence, decrease the risk of local recurrence. Surgery is the cornerstone of the treatment and inadequate local excision has been linked to local recurrence. Herein, we report a case of a 23-year-old female patient presenting with a large malignant phyllodes tumor of the breast treated by total mastectomy.

Case Presentation

Our case was a 23-year-old female patient Gravida 0 Para 0, previously healthy, with no family history of breast or ovarian cancer transferred to our institution to receive treatment for a right breast mass increasing in size over a period of 1 month associated with brownish and reddish discoloration of the skin. On physical examination of the right breast a grossly distorted architecture, with reddish-brown discoloration of the skin (Figure 1a, 1b) was observed

with no nipple retraction. On palpation of the diffusely hard breast, no nipple discharge and no palpable right axillary lymph nodes were observed. The contralateral breast (left) was normal looking, with no palpable masses, and no left axillary lymph nodes.

The patient underwent investigations with breast ultrasound showing an isodense oval mass measuring approximately 16x14x12cm in the right lower outer quadrant of her breast. Core biopsy and FNA done revealed fragments of myxoid and cellular stroma. These findings were not specific but could indicate a fibroepithelial lesion such as cellular fibroadenoma and low grade phyllodes tumor. MRI of the breasts showed a huge mass of the right breast occupying all quadrants, measuring 15x14.5 cm in its greatest axis with involvement of the pectoralis major muscle, with no evidence of suspicious axillary lymphadenopathy. Distant metastatic work-up prior to the surgical intervention with CT scan of chest abdomen and pelvis and a PET scan did not show distant metastasis. Hence, the patient was



Figure 3. Intra-operative image showing the surgical bed after total mastectomy



Figure 4. Surgical Specimen excised



Figure 5. Surgical Scar

scheduled for right total mastectomy.

The patient went through a right total mastectomy with partial excision of the pectoralis major where the tumor was involved (Figure 3, 4). Primary closure was possible without the need for autologous flap for closure. The final histopathologic result was consistent with features of malignant phyllodes tumor measuring 16.5x15x14cm, with clear negative surgical margins 1.6cm away from the tumor and the deepest surgical margin being 2mm away from the tumor. Post-surgical follow-up showed good wound healing, revealing no flap necrosis (Figure 5). She is on regular follow up and 18 months post-surgery is disease free with no evidence of local recurrence or distant recurrence by PET scan.

Discussion

Extensive studies have been done regarding the treatment of common breast malignancies. However, the optimal management for rare breast tumors including phyllodes tumor has not been well established. Due to its rarity, the current medical literature lacks a sound treatment plan, especially when it comes to the effect of chemotherapy or radiotherapy on managing malignant phyllodes tumor of the breast. Most management protocols depend on retrospective case series. Irrespective of their histologic classification, phyllodes tumors have an unpredictable clinical outcome. Age, tumor size, surgical approach, mitotic activity, stromal overgrowth and surgical margin have been found to be important factors in diagnosis and prediction of recurrence.⁶ Kaprisi et al. found that tumor size and surgical margins were predictive of local recurrence,⁷ which is very important for treatment of phyllodes tumors to be successful. A review of the literature reveals a local recurrence rate >40% for all pathological types.⁸ On the contrary, different studies showed a local recurrence rate of less than 20% depending on the tumor subtype. Furthermore, a delay in treatment increases the possibility of distant metastases. Metastasis usually occurs via the

bloodstream and lymphatic metastasis occurs rarely. The most frequently reported sites for metastases are the lung, soft tissue, bone, and pleura.⁹ The rate of metastases is 25%–31% for malignant and borderline tumors, whereas it is 4% for all types of phyllodes masses.¹⁰ Guidelines for cases of malignant phyllodes tumor by National Comprehensive Cancer Network include complete surgical resection with negative margins of at least 1 cm and preferably 1 to 2 cm due to the morphologic characteristic of the tumor. The presence of a pseudo capsule containing finger-like or pseudopodia-like projections around the phyllodes tumor tissue is likely, which, if not surgically excised, leads to a local recurrence. Therefore, the tumor with 1–2-cm width of normal tissue should be removed including these pseudopodia of tumors.^{11,12} Total mastectomy is in general not suggested due to poor cosmetic outcomes, unless negative margins cannot be conducted or if the tumor is too large to be completely excised. Adjuvant radiation therapy is debatable in preventing tumor recurrence. Radiation has been found to prevent recurrence of borderline and malignant phyllodes tumor after surgical interventions (breast-conserving surgery and mastectomy) with no impact on disease-free survival.¹³ Furthermore, adjuvant radiation therapy is suggested for recurrent and malignant phyllodes tumor. Patients with high risk for recurrence, including those with histological margin less than 5 mm may be candidates for radiotherapy.^{14, 15} On the other hand, routine adjuvant systemic therapy after initial excision is not recommended. Chemotherapy for locally recurrent tumor is contentious. Burton et al. reported effective palliation for patients suffering from metastatic disease with chemotherapy based on cisplatin and etoposide. As far as we know, this is the youngest reported case of malignant phyllodes tumor of the breast with this size. In our case, despite the large size of the tumor negative surgical margins of 1.6 cm were achieved by total mastectomy, with no recurrence at 18 months of follow-up. In our



review of the English medical literature we identified 16 cases of large malignant phyllodes tumors of the breast, above 10cm in largest dimension (Table 1), with a size range of 10 to 40cm, with a mean age of 44 years old. Table 1 shows that the most common site of metastasis is lungs 6/16

(37.5%), followed by brain 3/16 (18.7%). Hence, appropriate surgery is the cornerstone and the first step towards successful treatment of phyllodes tumor.

In conclusion, management of malignant

Table 1. Comparison the studies evaluated malignant phyllodes tumor

Author/year	Age	Pre-operative work-up	Size(cm)	Tumor Site	Type of Surgery	Adjuvant treatment	Recurrence
O.ECW*, K. CK, 2017 ¹⁶	53	Core biopsy inconclusive CT chest and abdomen: large tumor with extension to rib and pleura	40x38x16	Right	Radical mastectomy (tumor dissected off with right 6th rib and pectoralis major) (primary closure not possible)	None	1.All margins involved 2.Post-op PET scan : tumor infiltrating the 5th and 6th ribs 3. 6 weeks post-operative : re-admitted for pleural effusion → deceased due to pulmonary complications
A. Testori et al., 2015 ¹⁷	33	PET: positive ALN Skeletal & lung xray normal	40x30x10	Right	Modified radical mastectomy (1/9 LNs)	Chemotherapy (adriamycin and ifosfamide)/radiotherapy	Free of recurrence at 18 months follow up
R.Krishnamoorthy et al., 2014 ¹⁸	35	FNA: borderline phyllodes Core biopsy: biphasic neoplasm with malignant stromal component. CT brain and chest & US abdomen no metastasis	33x32x22	Right	Simple mastectomy	Chemotherapy (doxorubicin and ifosfamide) Radiotherapy	Free of recurrence at 1 year follow up
AP.Gregston et al., 2019 ¹⁹	32	Chest CT scan: large solid mass occupying the entire left breast	32x32x17	Left	Radical mastectomy with split thickness graft	6 cycles (AdriamycinVR, ifosfamide, and mesna) 4 cycles (gemcitabine and docetaxel)	2 months after surgery: 1.CT chest : pulmonary nodules •Biopsy : Lung metastasis •4 months after chemotherapy: progression of lung metastasis 2.Headache and grand mal seizure: brain metastasis (6x5x5 cm cerebral mass) underwent resection 3.Post-operatively dyspnea: intrathoracic progression of disease 4.Succumbed to disease 20 months after first diagnosis
N. Roberts, D.M. Runk, 2015 ²⁰	50	Core needle biopsy: malignant neoplasm with carcinomatous and sarcomatous elements, the cells showed focal marked pleomorphism (>10 mitoses/10HPF) with focal necrosis (ER/PR positive; HER2 negative) CT of chest positive for bilateral chest masses PET Scan: result not mentioned	31.5x15.6	Right	Simple mastectomy	Patient deceased before any therapy could be initiated	1.3 weeks post-op: complaint of headache Right parietal tumor – right craniotomy: metastatic malignant phyllodes tumor 2.7 weeks post-op •LR: 6 new masses were found, the largest measuring 6 cm and a mass was palpated in the right axilla: •FNA: enlarged pleomorphic cells, epithelioid cells – identical to original tumor. 3.Patient deceased before initiating treatment
I. Albalawi, 2018 ²¹	41	True cut biopsy malignant phyllodes tumor CT scan chest, abdomen and pelvis normal Isotopes scan normal	30x20x13	Left	Simple mastectomy	Radiotherapy	Free of recurrence 1 year after surgery
MA. Sbeih et al., 2015 ²²	41	Core biopsy: fascicular pseudoangiomatous stromal hyperplasia or phyllodes tumor Left ALN biopsy: chronic lymphadenitis without neoplastic cells CT chest and abdomen : no metastasis	20x20x25	Left	Simple mastectomy with left chest wall defect was covered temporarily with INTEGRA replaced with split thickness skin graft 5 weeks later	None	NA

M.Takenaka et al., 2011 ²³	57	Core bx low grade myofibroblastic sarcoma) CT suspicion of chest wall invasion MRI no ALN CEA/Ca 15-3 normal Excisional marked stromal hypercellularity without an epithelial component	20x14x9.7	Right	Radical mastectomy with pectoralis major excision+ axillary lymph nodes dissection (no positive ALNs)	None	No LR 17 months after surgery
Our Case, 2020	23	Core biopsy & FNA: fragments of myxoid and cellular stroma, may indicate a fibroepithelial lesion such as cellular fibroadenoma and low grade phyllodes tumor CT chest abdomen& pelvis negative for metastasis PET scan negative for metastasis	16.5x15x14	Right	Total mastectomy with partial excision of pectoralis major (with primary closure of defect)	None	No LR after 18 months follow-up PET scan: no distant metastasis
SH.Moon et al., 2019 ²⁴	48	Core needle biopsy: malignant phyllodes tumor CT chest metastatic LNs to Right axilla & multiple solitary pulmonary nodules on right lung FNA of LNs positive for metastasis PET scan	15.5x13.5x13	Right	Modified Radical Mastectomy + TRAM flap (1/32 ALNs)	Chemotherapy (doxorubicin and ifosfamide)	Free of LR at 3 years follow-up and free of distant metastasis (lung metastasis resolved)
T.Saito et al., 1998 ²⁵	54	NA	15x15	Right	Simple mastectomy	Radiotherapy (50Gy), Chemotherapy (Cyclophosphamide)	1. After 9 months Lung metastasis 2. Died of disease after 18 months of primary diagnosis
S. Khanal et al., 2018 ²⁶	37	Chest X-ray: negative US abdomen & pelvis negative True-cut: necrotic areas with fibrocollagenous tissue with proliferation of oval to elongated spindle cells showing mild degree of atypia	15x15	Left	Modified Radical Mastectomy	None	3 months' post-op: · No LR · Huge cystic nodule in the brain · Contrast enhanced computed tomography (CECT) of chest, abdomen and pelvis showed lesions in lower lobe of left lung of 3.9 × 3.6 cm of +40 (HU) in posterobasal segment and right adrenal gland of 5.8 × 5.1 cm of +30HU · 20 days' later patient succumbed to disease
J. Kaya, A. Betensley, 2013 ²⁷	48	NA	15	Left	Simple mastectomy	Radiotherapy	· 19 months post-op: left pleural effusion/left pleural mass · Patient deceased of disease 21 months after diagnosis
J.Alves et al., 2011 ²⁸	38	Core biopsy: biphasic neoplasm with malignant stromal component Chest X-ray & Abdomen ultrasound negative for distant metastasis	14x14x10	Right	Simple mastectomy without ALND	None	NA
A.Abdulkareem, 2019 ²⁹	37	Core biopsy: Epithelial and stromal components Epithelial is benign, the mesenchymal part showed spindle cell with increased cellularity (mitotic rate 5/10 HPF) no atypia or necrosis	13x11x7	Right	Partial mastectomy	Radio	1. Free margins but <1mm: re-excision margins 2. CT scan and bone scan negative for metastasis
M.Sani et al., 2008 ³⁰	44	FNA: benign breast lesion	12	Right	Modified radical mastectomy	None	Post-op CT chest and abdomen scan: no liver or lung mets Follow-up NA
S.Suzuki-Uematsu, 2010 ³¹	52	CT chest (bilateral lung nodules) Core needle biopsy (proliferation of atypical spindle-shaped cells with marked coagulation necrosis ,no epithelial components : sarcoma vs phyllodes)	10x10	Right	Simple mastectomy+ ALN excision	4 cycles (FEC therapy : cyclophosphamide, epirubicin, fluorouracil)	1. Lung metastasis increased in size (progressive disease) 2. Patient died 4 months post-surgery due to respiratory failure



phyllodes tumor remains debatable, especially when it comes to the role of adjuvant radiotherapy and chemotherapy. From a surgical perspective, whether phyllodes tumors should be considered as epithelial breast cancers or as soft tissue sarcoma is another debate. On the other hand, although challenging, accurate identification of phyllodes tumor initially may aid in decreasing the rate of local recurrence. Finally, due to rarity of such a disease, combining the observations of clinicians treating malignant phyllodes tumor in an international database will be beneficial to patients.

Acknowledgements

Special thanks to the General Surgery Department at Saint George Hospital University Medical Center and the National Breast Care Center at Sister Rosary Hospital

Ethical Consideration

A written informed consent was obtained from the patient to publish the case as well as any associated images

Conflict of Interest

None.

References

- Rowell MD, Perry RR, Hsiu JG, Barranco SC. Phyllodes tumors. *Am J Surg*. 1993;165(3):376-9.
- Bernstein L, Deapen D, Ross RK. The descriptive epidemiology of malignant cystosarcoma phyllodes tumors of the breast. *Cancer*. 1993;71(10):3020-4.
- Salvadori B, Cusumano F, Del Bo R, Delledonne V, Grassi M, et al. Surgical treatment of phyllodes tumors of the breast. *Cancer*. 1989;63(12):2532-6.
- Reinfuss M, Mitus J, Duda K, Stelmach A, Rys J, et al. The treatment and prognosis of patients with phyllodes tumor of the breast: an analysis of 170 cases. *Cancer*. 1996;77(5):910-6.
- Abusalem OT, Al-Masri A. Phyllodes Tumors of the Breast. *Mater Sociomed*. 2011;23(4):200-5.
- Briggs RM, Walters M, Rosenthal D. Cystosarcoma phylloides in adolescent female patients. *Am J Surg*. 1983;146(6):712-4.
- Mishra SP, Tiwary SK, Mishra M, Khanna AK. Phyllodes tumor of breast: a review article. *ISRN Surg*. 2013;2013:361469.
- Parker SJ, Harries SA. Phyllodes tumours. *Postgrad Med J*. 2001;77(909):428-35.
- Yagishita M, Nambu Y, Ishigaki M, Okada T, Yamanouchi K, et al. [Pulmonary metastatic malignant phyllodes tumor showing multiple thin walled cavities]. *Nihon Kokyuki Gakkai Zasshi*. 1999;37(1):61-6.
- Khosravi-Shahi P. Management of non metastatic phyllodes tumors of the breast: review of the literature. *Surg Oncol*. 2011;20(4):e143-8.
- Badwe RA, Kataria K, Srivastava A. Surgical Resection of Phyllodes Tumour: a Radical Approach as a Safeguard Against Local Recurrence. *Indian J Surg*. 2015;77(2):161-3.
- Kataria K, Dhar A, Ranjan P, Kumar A, Islam S, et al. Radical mastectomy SANS axillary lymph node dissection for large phyllodes tumor: a guarantee against recurrence. *Indian Journal of Surgery*. 2019;81(6):520-4.
- Zeng S, Zhang X, Yang D, Wang X, Ren G. Effects of adjuvant radiotherapy on borderline and malignant phyllodes tumors: A systematic review and meta-analysis. *Mol Clin Oncol*. 2015;3(3):663-71.
- Barth RJ, Wells WA, Mitchell SE, Cole BF. A prospective, multi-institutional study of adjuvant radiotherapy after resection of malignant phyllodes tumors. *Annals of surgical oncology*. 2009;16(8):2288-94.
- Chaney AW, Pollack A, McNeese MD, Zagars GK. Adjuvant radiotherapy for phyllodes tumor of breast. *Radiat Oncol Investig*. 1998;6(6):264-7.
- Ong E, Kong C. CASE REPORT: MASSIVE RUPTURED MALIGNANT PHYLLODES TUMOUR. *Journal of Health and Translational Medicine*. 2017;20(1):18-20.
- Testori A, Meroni S, Errico V, Travaglini R, Voulaz E, et al. Huge malignant phyllodes breast tumor: a real entity in a new era of early breast cancer. *World J Surg Oncol*. 2015;13:81.
- Krishnamoorthy R, Savasere T, Prabhuswamy VK, Babu R, Shivaswamy S. Giant malignant phyllodes tumour of breast. *Case Rep Oncol Med*. 2014;2014:956856.
- Gregston AP, Metter DM, Osborne CRC, Phippen J, Jr. Giant malignant phyllodes tumor with metastasis to the brain. *Proc (Bayl Univ Med Cent)*. 2019;32(1):116-8.
- Roberts N, Runk DM. Aggressive malignant phyllodes tumor. *Int J Surg Case Rep*. 2015;8C:161-5.
- Albalawi IA. A huge phyllodes tumor in the breast: a case report. *Electron Physician*. 2018;10(6):6951-5.
- Sbeih MA, Engdahl R, Landa M, Ojutiku O, Morrison N, et al. A giant phyllodes tumor causing ulceration and severe breast disfigurement: case report and review of giant phyllodes. *J Surg Case Rep*. 2015;2015(12).
- Takenaka M, Toh U, Otsuka H, Takahashi H, Iwakuma N, et al. Giant malignant phyllodes tumor: a case report. *Kurume Med J*. 2011;58(2):67-72.
- Moon SH, Jung JH, Lee J, Kim WW, Park HY, et al. Complete remission of giant malignant phyllodes tumor with lung metastasis: A case report. *Medicine (Baltimore)*. 2019;98(12):e16146.

- 2019;98(22):e15762.
25. Saito T, Kato K, Kawamori J, Yoshinobu T, Sato T, et al. Radiotherapy to pulmonary metastases of a malignant phyllodes tumor of the breast. *International Journal of Clinical Oncology*. 1998;3(5):337-41.
 26. Khanal S, Singh YP, Bhandari A, Sharma R. Malignant phyllodes tumor with metastases to lung, adrenal and brain: A rare case report. *Ann Med Surg (Lond)*. 2018;36:113-7.
 27. Kaya J, Betensley A. Is It Mass or Effusion: A Case of a Large Pleural Metastasis of Malignant Phyllodes Tumor of the Breast. *Chest*. 2013;144(4):484A.
 28. Souza JAd, Marques EF, Guatelli C, Girao DS, Queros T, et al. Malignant phyllodes tumor of the breast: case report. *Revista da Associação Médica Brasileira*. 2011;57(5):495-7.
 29. Abdulkareem AA. Malignant breast phyllodes: Literature review of management and case report. *Saudi Journal of Laparoscopy*. 2019; 4(1):1.
 30. Sani M, Leow V, Zaidi Z. Malignant phyllodes tumour: a case report. *The Internet Journal of Surgery*. 2008;21(2).
 31. Suzuki-Uematsu S, Shiraishi K, Ito T, Adachi N, Inage Y, et al. Malignant phyllodes tumor composed almost exclusively of a fibrosarcomatous component: a case report and review of malignant phyllodes tumors with metastases. *Breast cancer*. 2010;17(3):218-24.