

Recent Scenario of Male Breast Cancer in a Tertiary Care Centre

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Background: Breast cancer in males is a rare disease with an estimated incidence of 0.5-1% of all breast cancer cases. However recent data show that incidence is slowly rising but epidemiological data regarding Male Breast Cancer is little as compared to Female Breast Cancer due to the rarity of this disease. It is mostly seen in elderly men in the 6th or 7th decade of life. They usually present at an advanced stage with a poor prognosis as compared to FBC. This study is aimed to analyze the epidemiologic, and clinicopathological behavior of male breast cancer patients.

Methods: This work is the prospective work of 11 male breast cancer patients who were presented in department of General Surgery Rims, Ranchi during a period of 1 year. Data regarding clinical history, examination findings, family history, stage of the disease, histopathological examination and hormonal status were evaluated.

Results: Results showed that male breast cancer occurs later in life with median age being 58.5 years. The average diagnosis delay was 6.6 months. Out of eleven patients, eight of them had involvement of the left breast while the other three had involvement of right side of breast. Invasive ductal carcinoma was the histological type found in nine cases. Ten patients presented an advanced stage of T4 while one patient presented at T3. Hormone receptors were positive in 81.8% cases.

Conclusion: To conclude male breast cancer is a rare disease with multiple and varied risk factors. It is similar in clinical, histological and prognostic characteristic to breast cancer in women but also has varied characteristics. We suggest that for early diagnosis and enhanced prognosis, teaching self-breast examination in males, screening mammography and testing for familial predisposition shall be helpful in high risk groups. Therefore, there is a requirement for more free awareness public programmes to educate males about breast cancer and promote its early detection.

Introduction

Male breast cancer is a rare disease. It represents only 0.6% of all breast cancers and less than 1% of all causes of male cancer [1]. Male breast cancer is a relatively rare malignancy when compared to female breast cancer (FBC). The ratio of male breast carcinoma to female is 1:100 cases. Incidence is relatively low in men because of presence of less amount of breast tissue. It is also because of the difference in their hormonal environment when compared to female. In the past few decades concern about male breast carcinoma has increased, as the incidence has been rising rapidly. Review of Surveillance, Epidemiology and End Result data (SEER) indicates an [2,3] increased incidence of male breast cancer from 1.0 per 100,000 men in the late 1970s to 1.2 per 100,000 men from 2000 to 2004. Most male breast cancer occurs in elderly men. [4-6]. It is more commonly seen on left side with a ratio of 1.07:1 with respect to right side [7]. Diagnostic and treatment evidence for male breast cancer is limited. Currently, male breast cancer is treated

similar to female breast cancer [6,8].

Presentation

The most common presentations are painless palpable mass, skin ulceration, and nipple retraction or discharge in approximately 63 % of the cases, similar to women. As the males have relatively less breast tissue, nipple is mostly involved at early stages in most cases. The incidence of retraction of nipple is around 9%, for discharge it is 6% and ulceration being 6%. The mass is frequently localized to the subareolar region. It is seen less frequently in the upper outer quadrant. The left breast is involved more frequently than the right; 1% of the cases are bilateral. Others include Gynecomastia and Axillary lymphadenopathy (Figure 1-4).

Figure 1. Frontal View of Different Presentations of Patients. a, With nipple environment; b, Without nipple environment; c, Advanced stage cancer.

Figure 2. Frontal View of Specimen Taken after Modified Radical Mastectomy of Left Breast.

Figure 3. Frontal View of Scar Following Modified Radical Mastectomy Left Breast.

Figure 4. Left Breast Biopsy Positive for Invasive Ductal Carcinoma.

Diagnosis

Diagnosis of male breast cancer is mostly done by triple assessment similar to female breast cancer. Triple assessment which includes clinical assessment, mammography or ultrasonography, and core biopsy. Physical examination includes palpation of breast lump with axillary lymph nodes. The size of the tumour, it's consistency, boundary and mobility should be assessed. Male breast cancer is usually presents as hard mass, ill-defined margins and which has poor mobility, mostly fixed to breast tissue [9]. Ultrasound and trucut biopsy are routine preoperative examinations for breast cancer. Breast ultrasound shows usually a mass with unclear boundaries, incomplete capsule, and abundant vascular supply [10]. Mammography is difficult to carry out in male patients because of low breast tissue in males. Experienced mammography technicians are required for it if necessary. Mammography in male breast cancer shows high density mass with unclear boundaries and presence of burrs [7]. For most patients, enhanced magnetic resonance imaging is less available than ultrasound and mammography as it is costlier and more time consuming. Magnetic resonance imaging of male breast cancer shows irregular shape and enhanced signal of lesion. Based on all the above examination, we make the diagnosis of male breast cancer in clinic but the final diagnosis is made by histopathological examination.

Risk Factors

Male breast cancer etiology mostly includes the genetic or environmental risk factors. It is important to be mindful when discussing those risk factors in a case of male breast cancer. Majority of males who have been diagnosed with male breast cancer have no identifiable risk other than

increasing age (average age of diagnosis of 71 years) [4]. Risk of male breast cancer increases if they have first or second degree relative with breast cancer which is similar to cancer seen in women. Studies have shown that the affected first or second degree relative of either gender may increase the risk of breast cancer in either the males or females in the family.

Along with family history, increased risk for breast cancer in males is seen with BRCA mutation. It is mostly rare for a male to have a BRCA mutation, but those with BRCA2 carrier mutation have a 6% increased risk and BRCA1 carrier mutation have an increased risk of 4% for developing the disease [6]. Other genetic disorders which have been implicated in increasing the risk of male breast cancer includes Klinefelter syndrome (47, XXY), Cowden syndrome (PTEN tumour suppressor gene), Li-Fraumeni syndrome (TP53), Lynch syndrome (PALB2 and mismatch repair genes) [4][6].

The alterations in oestrogen to androgen ratios is important in etiology of male breast cancer and carries an important role. Same as female breast cancer oestrogen, which stimulates ductal development in breasts, has also been implicated as a potential risk factor for male breast cancer. Klinefelter syndrome as previously mentioned, increases the risk of male breast cancer by stimulating excess oestrogen. Other potential triggers include obesity, marijuana use, hepatic dysfunction, thyroid disease, and oestrogen-containing medications [4]. Similarly, a decrease in circulating androgens would result in elevations of the oestrogen, thus causing alteration in oestrogen and androgen ratios, and predisposing to male breast cancer. This alteration is mostly seen in conditions such as cryptorchidism, orchitis, and orchiectomy [8].

Lastly, environmental factors have also been playing an important role for increasing the risk of male breast cancer. In males' previous radiation therapy has been noted as a potential risk factor like females. Radiation therapy increases the risk of male breast cancer 7 times. In males generally radiotherapy has been used to treat gynecomastia [2]. In addition to all this, certain occupations have also been implicated to increase the risk of MBC, including steel and rolling mills, as well as occupations involved with organic solvents such as trichloroethylene [3].

Aims and Objectives

- To discuss the epidemiological, clinical, and histological features of male breast cancer.
- To discuss the therapeutic management as well as the survival rates and factors of this disease.

Materials and Methods

This work is prospective work of 11 male breast cancer patients who presented in department of general surgery Rims Ranchi during a period of 1 year. Data regarding clinical history, examination findings, family history, stage of the disease, histopathological examination and hormonal status were evaluated.

Inclusion Criteria:

a. All male patients admitted in surgical ward through OPD/ Emergency with breast lump or ulcer.

b. Male patients aged more than 18 years. Exclusion Criteria:

a. Patients who succumbed to death during the course of treatment.

b. Study patients who had no follow-up after initial diagnosis.

Variables Studied:

- Age at diagnosis
- Family history
- Tumour size (in cm)
- Grade
- Stage of the tumour
- Histological subtype of the tumour
- Axillary lymph node status
- Average delay in diagnosis
- Oestrogen receptor (ER) expression,
- Treatment

Results

In my study, Male breast cancers occurred later in life with late stage and more ER PR receptor-positive tumours. The Median age being 58.54 years. Family history was found positive in one case. The average Mean diameter of lump was 8.77 cm (range: 3-20 cm) and tumor was predominantly on the left side (72.72%). The Average diagnosis delay for patients was 6.66 months. According to TNM classification 10 patients presented at T4 stage (90.90%) and one patient presented at T3. Axillary nodal involvement was seen in 72.72% of cases. Three patient was Node negative (27.27%) and N1 was seen in 6 cases (54.54%), while N2 was seen in 2 case (18.18%). Histopathological finding was infiltrating ductal carcinoma in 9 cases. (81.8%), One case was ductal carcinoma in situ and one was spindle cell carcinoma. Hormone receptors (ER/PR) was positive in 81.81% cases. Rest patients were not financially supportive for the test. Neoadjuvant chemotherapy with CAF regimen followed by Modified radical mastectomy followed by adjuvant Chemotherapy was given to 7 patients. Neoadjuvant chemotherapy with CAF regimen followed by Modified radical mastectomy and axillary lymph node sampling followed by adjuvant Chemotherapy was given to 2 patients One patient underwent mastectomy and one underwent WLE (Table 1).

		Case 1	Case 2	Case 3	Case 4	Case 5	Case 6	Case 7	Case 8	Case 9	Case 10	Case 11
1	Age at diagnosis	68 years	70 years	58 years	60 years	56 years	50 years	57 years	74 years	62 years	20 years	69 years
2	Family history	Negative	Negative	Positive	Negative	Negative	Negative	Negative	Negative	negative	Positive	Negative
3	Mode of presentation	Swelling +	Swelling +	Swelling -	Swelling +	Swelling -	Swelling -	Swelling +	Swelling +	Swelling -	Swelling +	Swelling +
		Ulceration +	Ulceration +	Ulceration +	Ulceration +	Ulceration +	Ulceration +	Ulceration +	Ulceration +	Ulceration +	Ulceration -	Ulceration +
		Pain -	Pain +	Pain -	Pain -	Pain +	Pain -	Pain -	Pain -	Pain -	Pain -	Pain -
		Nipple distortion or discharge +	Nipple distortion or discharge +	Nipple distortion or discharge +	Nipple distortion or discharge +	Nipple distortion or discharge +	Nipple distortion or discharge +	Nipple distortion or discharge +	Nipple distortion or discharge -	Nipple distortion or discharge +	Nipple distortion or discharge -	Nipple distortion or discharge +
4	Average	14	18	5	6	10	7	4	5months	3months	2months	6

	diagnosis delay	months	months	months	months	months	months	months	months	months	months	months
5	Tm size (in cm)	14*18*6	20*10*8	4*4*3.5	7*5*3	16*14*8	3*2*2	7*6*4	6*5*3	4*3*1.5	3.5*2*1.5	8*6*4
6	Side of tumor	Left	Left	Left	Left	Left	Left	Right	Left	Right	Right	left
7	Axillary LN status	Positive	Positive	Negative	Positive	Positive	Positive	Positive	Positive	Positive	negative	negative
8	TNM Stage	T4bN1M0	T4bN2aM0	T4bN0M0	T4BN1M0	T4bN1M0	T4bN1M0	T4BN1M1	T4BN2aM0	T4BN1M0	T3N0M0	T4BN0M0
9	Histological subtype	Ductal	Ductal	Ductal	Ductal	Ductal	Ductal	Ductal	Ductal	Ductal	Ductal carcinoma in situ	Spindle cell carcinoma
10	Grade	Poorly differentiated	Poorly differentiated	Mod differentiated	Mod differentiated	Poorly differentiated	Well differentiated	Mod differentiated	Mod differentiated	Mod differentiated	Well differentiated	Well differentiated
11	Stage	IIIB	IIIB	IIIB	IIIB	IIIB	IIIB	IV	IIIB	IIIB	IIIA	IIIB
12	ER/PR status	ER +	ER +	ER +	ER +	ER +	ER +	ER +	ER +	ER +	Not done	Not Done
		PR -	PR +	PR +	PR +	PR -	PR +	PR -	PR +	PR +		
		Her 2 neu -	Her 2 neu -	Her 2 neu -	Her 2 neu -	Her 2 neu -	Her 2 neu -	Her 2 neu -	Her 2 neu -	Her 2 neu -		
13	Treatment	Left MRM	Left MRM	Left MRM	Left MRM	Left MRM	Left MRM	Right MRM	Left MRM	Right MRM	Mastectomy	WLE
14	HPE	Margin not involved	Margin not involved	Margin not involved	Margin not involved	Margin not involved	Margin not involved	Margin not involved	Margin not involved	Margin not involved	Margin not involved	Margin not involved
		Node involvement present	Node involvement present	Node involvement absent	Node involvement present	Node involvement present	Node involvement present	Node involvement present	Node involvement present	Node involvement present	Node involvement absent	Node involvement absent
		Lymphovascular invasion present	Lymphovascular invasion present	Lymphovascular invasion absent	Lymphovascular invasion absent	Lymphovascular invasion present	Lymphovascular invasion absent	Lymphovascular invasion absent	Lymphovascular invasion absent	Lymphovascular invasion absent	Lymphovascular invasion absent	Lymphovascular invasion absent

Table 1. Clinicoepidemiological and Pathological Profile of Male Breast Cancer Patients.

Discussion

The average age of our patients (58.54 years) which is almost similar to study done by A. Sanguinetti et al. (2016) [10] which is 62 years and Chao li and Xujung li et al (2022) which is 60 years. In study done by Majdouline El Fouhi et al (2022) [11] it was slightly higher, that is 67.7 years. Collective reviews have shown predilection for the left side in a ratio of 1.07:1. In our series also more patients had lump on the left side. (72.72%). The average diagnosis delay of our patients was 6.66 months which is slightly lower than that seen in other series, In studies Figure 1. Frontal View of Different Presentations of Patients. a, With nipple environment; b, Without nipple environment; c, Advanced stage cancer done by A. Sanguinetti et al. (2016) [10] it was 16 months and study done by Majdouline El Fouhi et al (2022) [11] it was 15.7 months. 81.8% cases of male breast tumors have turned out to be invasive ductal carcinomas in our study, expressing high levels of hormone receptors. In studies done by A. Sanguinetti et al. (2016) [10], Chao li and Xujung li et al (2022) and Majdouline El Fouhi et al (2022) (11), 95% (45/47), 80.7% (21/26), 92% (23/25) patients had Infiltrating ductal carcinoma (IDC) respectively. All these studies show that infiltrating ductal carcinoma is the most predominant subtype with an incidence ranging from 64-93%. Axillary lymph node involvement is very common and we report a rate of 72.72 % of axillary nodal involvement which is almost similar to study done by A. Sanguinetti et al. (2016) [10]: 79.4% (39/47) and Majdouline El Fouhi et al (2022) [11]: 80% (20/25) but it differed with study done by

Chao li and Xujung li et al (2022) which showed 53.84% (14/26) axillary lymph node involvement. Breast cancer in males should be treated with the same strategy in women. The most common surgical procedure is modified radical mastectomy with axillary node dissection. In our series, modified radical mastectomy was the common surgical procedure combined with adjuvant chemotherapy. Frequently used chemotherapy regimens were cyclophosphamide adriamycin fluorouracil (CAF). Majority of our patients have received the same. In My study Hormonal receptors (both ER and PR) were positive in 54.54% of case and ER was positive in 27.27% cases. In studies done by Sanguinetti et al. (2016) [10], 67% (31/47) were ER/ PR positive. Chao li and Xujung li et al (2022), 96% (25/26) were Both ER+ and PR+ and in study done by Majdouline El Fouhi et al (2022) [11], 83% (15/26) were ER+/PR+. Patients who showed positivity of ER/PR in MBC (83%) had received hormone therapy (tamoxifen). Tamoxifen has been shown to lead to increased survival rates in women with hormone-sensitive disease and today is generally considered the standard adjuvant treatment for male breast cancer hormone-dependent. Also, hormonal therapy has been proven to help in metastatic disease in females and males.

In conclusion, hence, concluding from the above discussion, Male breast carcinoma presents later in life (age>60years) and it presents in advance stage of carcinoma(T4). Most common histopathological subtype is infiltrating ductal carcinoma. Most of the patients are receptor positive (ER/PR). Therefore Concerted efforts including education of public and health professionals, in order to make earlier diagnosis and thereby improve prognosis. Efforts should be made to improve outcomes, an interprofessional team approach that consists of an oncologist, surgeon, radiation therapist, dietitian, and mental health counselor is recommended. Earlier diagnosis and wide use of adjuvant treatments (RT/HT/ CT) widely decreased local recurrences and increased survival rates in male breast cancer .

Acknowledgments

We acknowledge the help extended by the Department of General Surgery, Rajendra Institute of Medical Sciences Ranchi, India.

Conflicts of Interest

The authors have no conflicts of interest to declare.

Funding statement

There was no source of funding for this research

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