

**Male Breast Disease** 

Home Study Continuing Education Course

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# EDUCATIONAL REVIEW

# Imaging findings and classification of the common and uncommon male breast diseases

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#### Abstract

Male breast hosts various pathological conditions just like "female breast." However, histo-anatomical diversities with female breast lead to many differences regarding the frequency and presentation of diseases, the radiologic appearance of lesions, the diagnostic algorithm, and malignity features.

Radiological modalities may play an important role in evaluating male breast lesions. Although some imaging findings are non-specific, having knowledge of certain imaging characteristics and radiologic patterns is the key to reduce the number of differential diagnoses or to reach an accurate diagnosis.

Male breast imaging is mostly based on physical examination and is required for the complaints of palpable mass, breast enlargement, tenderness, nipple discharge, and nipple-skin changes. The majority of the male breast lumps are benign and the most common reason is gynecomastia. Although it is difficult to exclude malignancy in some cases, gynecomastia often has distinguishable imaging features. Pseudogynecomastia is another differential diagnosis that may be confused with gynecomastia. The distinction is important for the treatment plan.

Apart from gynecomastia, other male breast lesions form a highly heterogeneous group and can be classified based on "Tissue origin," "Histopathological type and behavior," and "Radiologic features" for both simplification and comprehensive understanding.

This article mainly focuses on emphasizing the results of basic histo-anatomical differences of male and female breasts, classifying male breast lesions, covering the spectrum of male breast diseases, and assisting radiologists in recognizing the imaging findings, in interpreting them through a holistic approach, in making a differential diagnosis, and in being a part of proper patient management.

Keywords: Male breast lesion, Classification, Ultrasound, Mammography, Imaging findings

#### **Key points**

- Male and female breasts have certain histoanatomical differences that can affect the spectrum of diseases.
- The majority of male breast lumps are benign and the most common reason is gynecomastia. Pseudogynecomastia and malignancy are the most common differential diagnoses.

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- Male breast imaging is mostly based on "physical examination," and radiological modalities can play an important role.
- Imaging of the male breast has many aspects regarding not only medical and scientific but also social concerns that can lead to poor patient compliance and follow-up losses.
- Male breast lesions form a highly heterogeneous group and could be classified based on various parameters for comprehensive understanding.

#### Introduction

The male breast is located between the 2nd-6th ribs craniocaudally and the midaxillary line-sternum

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lateromedially, like female breast. Although they share similar locations, there are several differences between them, which concern the developmental process and histo-anatomical structures [1].

The mammary glands of both sexes are identical at birth. During peripubertal period, in the female breast, ductal proliferation, branching, and growth are stimulated due to estrogen. As for stromal development and terminal ductal-lobular unit (TDLU) maturation, they are seen due to progesterone. However, involution and ductal atrophy occur in the male breast due to significant increase in testosterone levels [2].

In contrast to the female breast, Cooper ligaments are absent, ductal system is involuted, TDLU development is rare, stromal system is smaller in size and pectoralis muscles are more prominent in the male breast [3] (Fig. 1).

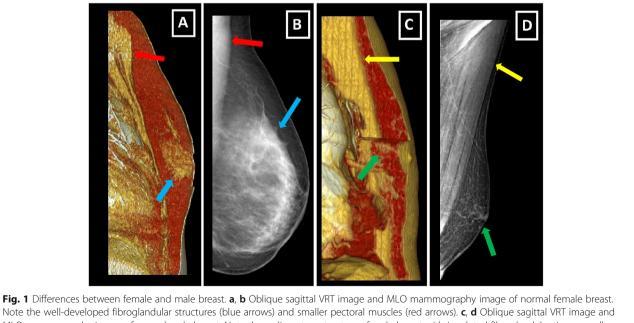
Involution of ductal system causes decreased ductal branching, so ductal malignancies are rarely encountered in males and are located in close proximity to nippleareolar complex when they exist. Breast cancer incidence is low in the male population, so there is no need for routine screening. Due to delay in diagnosis, male breast cancer is detected at a more advanced stage when compared with female breast cancer. At the time of diagnosis, axillary lymphadenopathy (LAP) involvement is at nearly 50%, which is more frequent than female breast cancer, and secondary signs, such as skin thickening, ulceration, and increased trabeculations, appear earlier. Moreover, microcalcifications, which may show ductal involvement, are seen less likely in male breast cancers, probably because of involuted ductal structure. Other than that, fibroepithelial (biphasic, i.e., fibroadenoma, phylloides tumor) and lobular (invasive lobular carcinoma (ILC), lobular carcinoma in situ (LCIS), fibrocystic changes, adenosis) pathologies are uncommon due to rare development of TDLU [2–6] (Fig. 2).

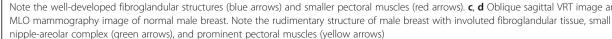
Male breast as a rudimentary structure consists of subcutaneous adipose tissue, remnant ductal tissue, and small nipple-areolar complex (Fig. 1). Keeping structure and components of the male breast in mind facilitates radiologic assessment. Mammography, ultrasound (US), and rarely magnetic resonance imaging (MRI) are the main imaging modalities (Fig. 3).

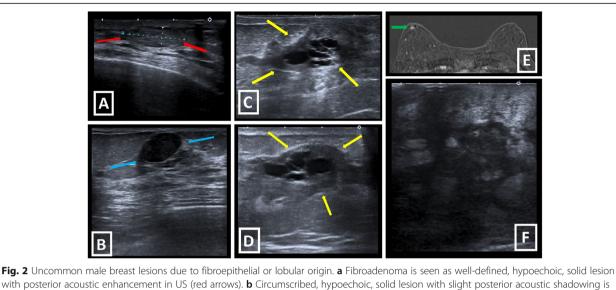
In mammography, a normal male breast is composed of homogenously radiolucent fat tissue and prominent radio-opaque pectoralis muscle. Small volume of male breast may cause technical difficulties. Moreover, prominent pectoralis muscle may obscure suspicious lesions [1, 4, 6, 7].

US imaging of the male breast has similar characteristics to that of the female breast. Isoechoic fat lobules corresponding to subcutaneous adipose tissue is a major finding of normal male breast seen in US. US can be very helpful for evaluating nipple-mass relationship and axillary nodal involvement. In case of suspicious lesions, vascularity assessment can be done using the Doppler mode. Sonoelastography may be used as auxiliary modality, although not on a routine basis [6–9].

MRI of male breast has limited use. According to literature, it can be used for evaluation of chest wall







with posterior acoustic enhancement in US (red arrows). **b** Circumscribed, hypoechoic, solid lesion with slight posterior acoustic shadowing is diagnosed as "Adenosis" after biopsy (blue arrows). **c**, **d** Fibrocystic changes with apocrine metaplasia. US shows multiseptated cystic structures within retroareolar heterogeneous hypoechoic area in a 59-year-old male patient with palpable mass (yellow arrows). **e** After detecting right retroareolar ductal ectasia with peripheral hypoechoic heterogeneous area in US (not shown), subtraction image of dynamic MRI of 72-year-old male patient shows retroareolar well-defined enhancing focus which corresponds to the defined lesion. Biopsy reveals the diagnosis of "Fibroadenomatoid changes". **f** Large, heterogeneous, ill-defined, hypoechoic mass with subcutaneous edema and pectoral muscle invasion (not shown) is diagnosed as LC

involvement of malignity, post-operative residual disease, and neoadjuvant chemotherapy response [10, 11].

Imaging of the male breast has an impact not only on medical and the scientific field but also on social issues. It could turn out to be somewhat problematic due to the social prejudices it is often associated with this. Social stigma related to male patient undergoing mammography or breast US within breast imaging centers, which are mostly used by female patients, may cause poor patient compliance and decreases in follow-up attendance [12].

There is no widely accepted consensus for diagnostic imaging algorithm of the male breast lesions. According to Mustapha et al., mammography is the first-line suggested imaging modality in case of suspicious physical examination. US is the next recommended step when mammography findings are inconclusive [13]. On the other hand, ACR appropriateness criteria suggest an age limit of 25 years old for first-line imaging modality. US is recommended as first-line imaging below 25 years old, whereas mammography is suggested after 25 years old. According to ACR, further imaging is not required when clinical findings are consistent with gynecomastia or pseudogynecomastia. However, mammography is proposed as a first step, if suspicion of cancer is high based on clinical findings [14].

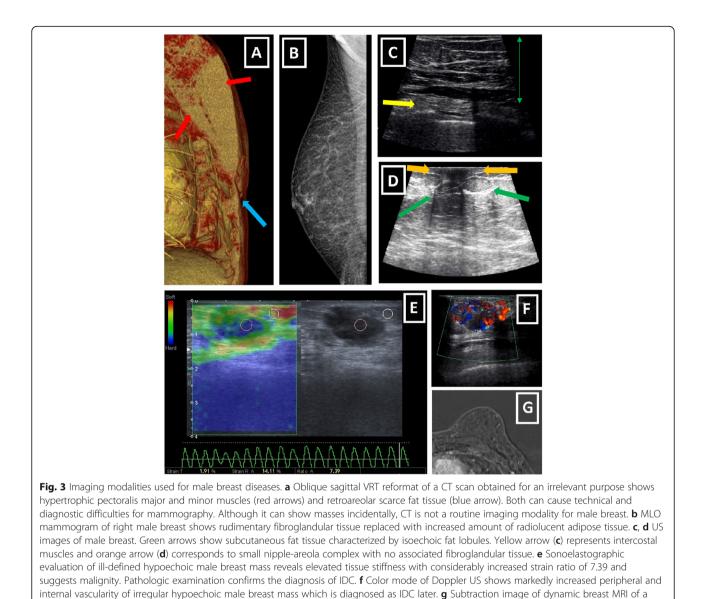
Biopsy is required for pathological diagnosis when imaging modalities are equivocal. For male breast, US guided biopsy is preferred, as stereotactic biopsies are technically difficult due to smaller breast sizes [3, 6, 15].

#### Gynecomastia, types, and differential diagnosis

The majority of male breast lumps are benign, and the most common reason is gynecomastia. Gynecomastia is formed by the proliferation of ductal and stromal elements, mostly seen during pubertal period or senescence, either presented as unilateral or bilateral breast mass along with breast enlargement and/or focal pain. Etiology of gynecomastia contains a wide-range spectrum including physiologic, endocrinologic, metabolic, neoplastic, and drug-induced causes. There are three characteristic patterns related to gynecomastia which are nodular, dendritic, and diffuse glandular forms [3, 6, 7] (Table 1, Fig. 4).

The nodular form is characterized by prominent ductal hyperplasia and cellular/proliferative stroma. Generally, it is the sign of the early (florid) phase corresponding to a duration of less than 1 year. Nodular gynecomastia is accepted as a reversible pathology if underlying cause(s) is eliminated. It is seen as nodular or fan-shaped subareolar opacity in mammography and disk-shaped, hypervascular, hypoechoic subareolar tissue in US [6, 7, 15].

The dendritic form corresponds to periductal stromal fibrosis and minimal ductal hyperplasia. It is known as the late (quiescent) phase of a duration exceeding 1 year. It is irreversible due to chronic changes and fibrosis. Flame-shaped subareolar opacity which may extend toward upper-outer quadrants is an expected mammographic finding. Subareolar serpiginous tissue with "star-



shaped" borders and "spider leg" appearance are the fibr main characteristic findings of US [3, 6, 7, 15, 16].

male patient with known pseudogynecomastia shows enlarged left male breast without apparent enhancing lesion

The diffuse form has the properties of both early and late phases. It is typically related to high-dose estrogen therapy or exogenous hormone usage. Transgender male breasts usually show the diffuse pattern. Heterogeneously enlarged breast tissue with female breast appearance is seen in mammography. The main anatomic difference between diffuse gynecomastia and female breast is the absence of Cooper ligaments. In US, increased breast volume and echogenicity are present which may be falsely interpreted as malignity [3, 6, 7, 15, 16].

The most common differential diagnoses of gynecomastia are pseudogynecomastia and malignancy. Pseudogynecomastia refers to diffuse adipose tissue proliferation without fibroglandular development or discrete mass in male breast, which can be unilateral or bilateral. It is mostly seen in overweight–obese individual or with those with Neurofibromatosis type 1 (NF-1) history. Differentiation from gynecomastia could be important for the treatment plan. Liposuction alone is mostly sufficient for the treatment of pseudogynecomastia. On the other hand, surgical excision of glandular tissue can be required for a satisfactory treatment of gynecomastia [3, 4, 6, 17] (Table 2, Figs. 3b and 4d).

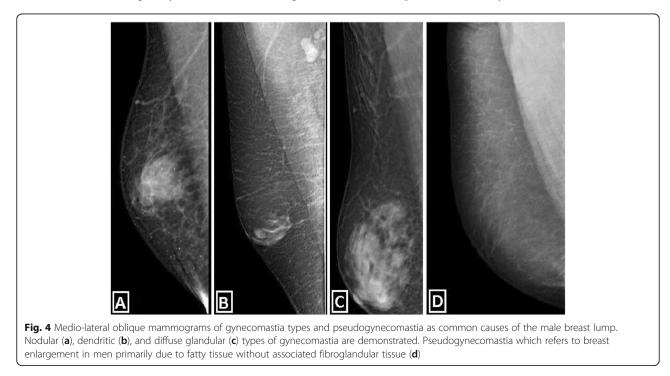
The distinction of malignancy and gynecomastia is critical because further intervention can be avoided when imaging findings are consistent with gynecomastia. Unfortunately this distinction is not always clear-cut. Although gynecomastia often has characteristic clinical

#### Table 1 Gynecomastia types

Gynecomastia Types	Nodular form	Dendritic form	Diffuse form
Phase	Early (florid)	Late (quiescent)	Both
Duration	<1 year	>1 year	-
Reversibility	Reversible (if stimulus is eliminated)	Irreversible	
Cause	Many	Many	Typically related with high-dose estrogen theraphy or exogenous hormon usage.
Mammographic (MMG) findings	Nodular or fan-shaped subareolar density.	Flame-shaped subareolar density. Could be extend toward upper- outer quadrants.	Both nodular and dendritic pattern, increased breast density, "female breast" appearance. Heterogenously enlarged breast tissue.
Ultrasonographic (US) findings	Disc-shaped hypervascular hypoechoic subareolar tissue.	Subareolar serpiginous hypoechoic tissue. "Star shaped" borders. "Spider legs" appearance.	Both nodular and dendritic patterns. Increased breast volume and echogenicity. Could mimic malignancy.
Histopathological findings	Epithelial hyperplasia. Cellular and proliferative stroma.	Minimal ductal hyperplasia and periductal stromal fibrosis.	Both patterns.

and radiological features which help distinguish it from malignancy, it may be difficult to exclude malignancy in some cases (Table 2, Fig. 5).

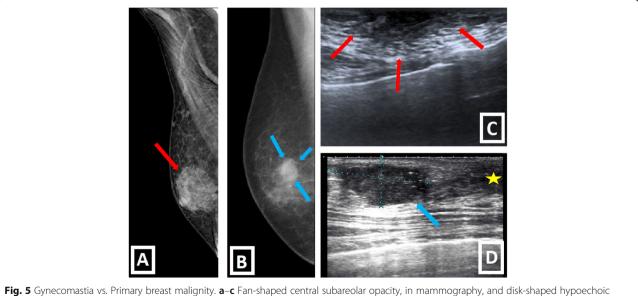
While gynecomastia has the abovementioned risk factors, the risk factors of malignancy are different such as genetic background (family history, BRCA2 mutation, Klinefelter syndrome), liver disease, prior radiotherapy history, endocrinologic, or drug-related causes. Gynecomastia has bimodal age distribution involving the peripubertal period and the period after 50 years. On the other hand,

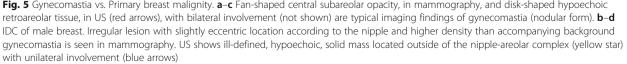


#### Table 2 Differential diagnosis of gynecomastia

Differential diagnoses	Pseudogynecomastia	Gynecomastia	Malignancy
Etiology	Mostly seen in overweight / obese individuals or with NF- 1 history.	Could be physiologic, drug- induced, endocrinologic, paraneoplastic, secondary to liver and kidney disease or idiopathic.	Risk factors: Genetic background (Family history, BRCA 2, Klinefelter syndrome), endocrinologic causes, drug-induced (hyperestrogenism), secondary to prior RT,liver disease
Age of presentation	Any	Peripubertal period and >50 y.o.a. (Bimodal distribution)	Generally after 60 y.o.a
Physical examination	Breast enlargement without a discrete mass.	Soft, tender, mobile, subareolar lesion.	Soft or firm, mobile or non-mobile, mostly non-tender lesion.
Laterality	Unilateral or bilateral	Mostly bilateral	Generally unilateral
Location (according to the nipple)	Diffuse proliferation of fat tissue without fibroglandular development or discrete mass.	Central. May extend toward upper-outer quadrants (But never only eccentric).	Could be eccentric (Important for differentiation from gynecomastia).
Secondary signs	Not expected	Not expected	Axillary LAP, increased trabeculation, skin thickening, ulceration, nipple retraction could be present.
Discrete masses, cystic components or complex appearance in US	Not expected	Not expected	Suspicious for malignancy

malignancy usually occurs after 60 years of age. Bilateral involvement is usually expected in gynecomastia while unilateral involvement is usually expected in malignancy. Central involvement is a rule for gynecomastia. Although extension to the upper-outer quadrants may be encountered, pure eccentric lesions are not expected in gynecomastia. In the malignancy, lesions can be eccentric, and it is important for differentiation from gynecomastia. Gynecomastia mostly present as soft, tender, mobile subareolar lesion. On the other hand, malignancy can be encountered as soft or firm,





mobile or non-mobile, and mostly non-tender lesion. Accompanying secondary signs including skin thickening, ulceration, nipple retraction, increased breast trabeculation, and axillary LAPs, favors malignancy. Discrete masses, cystic components, and complex sonographic appearances also suggest malignant pathology [3, 6, 7, 15].

#### Classification and spectrum of disease

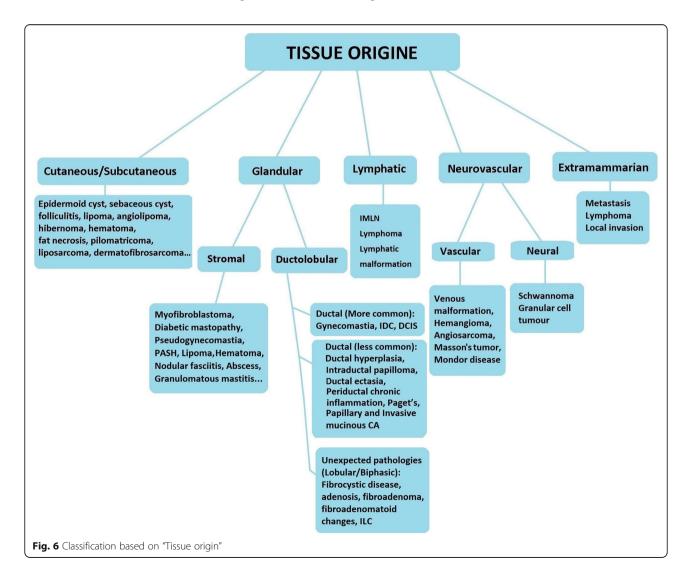
Apart from gynecomastia, male breast lesions form a highly heterogenous spectrum which will be discussed elaborately. They can be grouped based on various parameters, such as "Tissue origin," "Histopathological type and behavior," and "Radiologic features," for simplification, comprehensive understanding, and reasonable radio-pathologic integration.

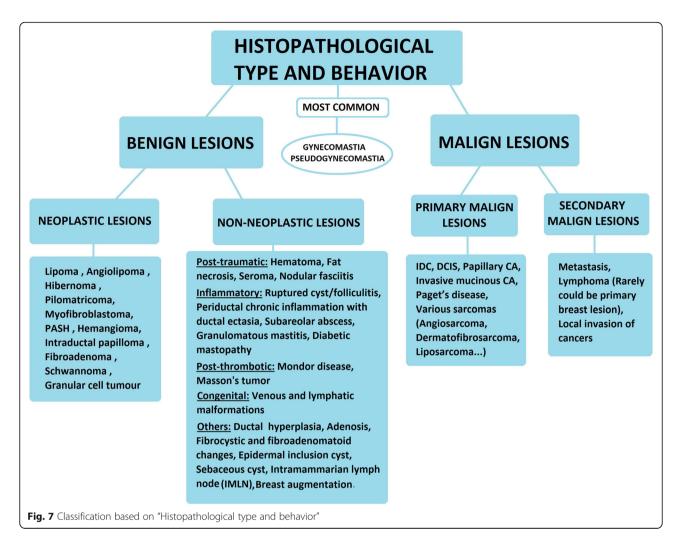
#### Tissue origin

Regarding tissue origin, male breast lesions may develop from cutaneous/subcutaneous tissues, glandular/stromal breast tissues, neurovascular tissues, lymphatic tissues, and extramammarian lesions (Fig. 6). Breast lesions from extramammarian origin are further classified, mainly, as solid organ metastases (prostate, lung, gastric cancers), and hematological malignancy involvement (lymphoma, leukemia, plasmacytoma). Local invasion of malignancies in close proximity to the breast, such as lung cancer, is also a probability [1, 15, 18].

#### Histopathological type and behavior

According to histopathological type and behavior we can divide male breast lesions into two main categories as "benign" and "malign." Benign lesions are subdivided into two parts as "neoplastic" and "non-neoplastic." Non-neoplastic benign lesions are further classified according to their etiologies. Malign lesions are classified as "primary" and "secondary" according to their origins (Fig. 7) [3, 6, 7].





#### Radiological features

We can classify male breast lesions according to their known radiological features including margins, internal structure, presence of calcification and vascularity, hyperechogenicity, and posterior acoustic features. In general, benign neoplastic lesions have well-defined margins. On the other hand, not only most of the malign tumors but also the post-traumatic and inflammatory benign nonneoplastic lesions have ill-defined/irregular margins. Additionally, increased vascularity is more common in these groups. Increased echogenicity is less common in malign lesions. The presence of calcification and cystic/tubular internal structures can be important clues for narrowing the differential diagnoses. Furthermore, having knowledge of malignity-mimicker lesions and their clinical presentation can be helpful to the radiologist for being a part of proper patient management and facilitating diagnostic work-up. As a result, preventing unnecessary intervention and relieving patient anxiety can be accomplished with the knowledge of pitfalls which mimic malignity [6, 15]. For example, imaging appearances of acute hematoma/fat necrosis may mimic malignancy, but history of trauma to the lesion site is an important reason for recommending follow-up, and avoiding immediate biopsy. The comprehensive list of disease subgroups with their radiological features are shown in Fig. 8.

#### Spectrum of disease

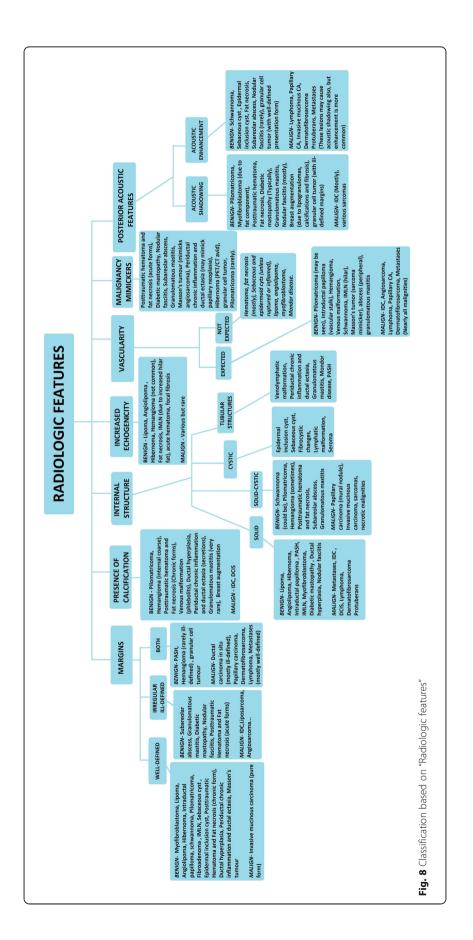
#### Benign neoplastic breast lesions

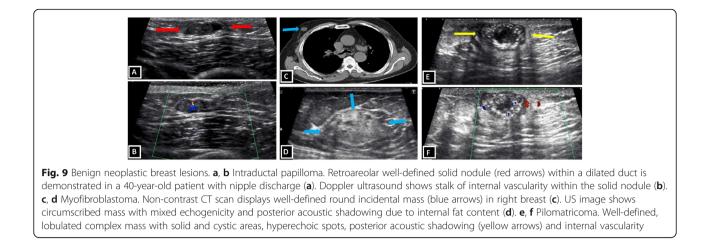
#### Lipoma

Lipoma is the second most common benign male breast lesion which is composed of mature fat cells and is presented as asymptomatic or soft, non-tender palpable mass on physical examination. Radiologically, lipoma is seen as well-defined, subtle encapsulated, radiolucent lesion in mammography and avascular, oval-shaped, parallelly oriented, mild hyperechoic or iso-hypoechoic mass with thin echogenic capsule in US [3, 6, 15, 16].

#### Angiolipoma

Angiolipoma, which is a benign tumor formed by mature adipocytes, thin-walled vessels, and fibrin thrombi,





is located typically anterior to the pectoralis fascia within fat tissue. Usual physical examination findings are painless, palpable, generally firm, mobile, and non-tender mass. Typical radiological findings are mass which is composed of mixed fat and soft-tissue densities in mammography and homogenous echogenic mass in US. Moreover, angiolipoma may not have any definite abnormality, in mammography. Differential diagnoses include acute hematoma, focal fibrosis, hemangioma, and malignancy [6, 19].

#### Hibernoma

It is slow-growing benign neoplastic lesion arising from brown adipose tissue. The physical examination findings are frequently mobile, painless, palpable, slow-growing mass. Hibernoma is an analogous of lipoma considering imaging findings. Sometimes, hibernoma may mimic malign lesions in different modalities. For example, in positron emission tomography-computed tomography (PET-CT) with fluorodeoxyglucose (FDG)-avidity, it may be impossible to distinguish it from well-differentiated liposarcomas reliably [20, 21].

#### Pseudoangiomatous stromal hyperplasia (PASH)

PASH is an uncommon stromal neoplasm composed of myofibroblasts with glandular hyperplasia within dense collagenous stroma. It is one of the benign conditions associated with gynecomastia, like as intraductal papilloma. PASH has two known forms: nodular mass-like and diffuse forms. Beside this, it could be seen incidentally within another lesion. Clinically, this benign lesion may be either asymptomatic or may present with focal palpable mass, especially in nodular form. Mammographic findings are "totally or partially circumscribed noncalcified breast mass" or "linear densities radiating from nipple similar to dendritic gynecomastia." Sonographic imaging findings are "solid circumscribed, parallelly-oriented, hypoechoic mass with/without heterogeneity" or "retroareolar hypoechoic area with projecting tubular structures like gynecomastia" [7, 15, 16, 22].

#### Intraductal papilloma

It is a benign neoplasm of the intraductal epithelium with fibrovascular core. It may present with nipple discharge or palpable, painless, mobile, subareolar mass, on physical examination. Intraductal papilloma is, also, one of the benign conditions associated with gynecomastia. Imaging findings are discrete, dense mass in subareolar region with or without gynecomastia in mammography and subareolar, eccentric, elongated, well-defined, hypoechoic mass within dilated cystic ducts in US. "Stalk" of internal vascularity can be seen in Doppler US imaging. The differentiation of intraductal papilloma from papillary carcinoma is not possible with imaging alone; that is why the pathologic correlation is needed [6, 15, 16] (Fig. 9a, b).

#### Myofibroblastoma

Myofibroblastoma is an uncommon benign mesenchymal neoplastic lesion of the breast. It is more frequent in males than females and it affects mostly adult male population. Myofibroblastoma is seen as a mobile, well-defined, solid lump, on physical examination. Imaging findings identified in mammography are well-defined, encapsulated, heterogeneous tumor without microcalcifications. Sonographic findings are well-demarcated tumor, mixed echo pattern, and acoustic attenuation, probably, due to fat component [7, 23] (Fig. 9c, d).

#### Pilomatricoma

It is rare benign neoplastic skin tumor originating from piliferous follicles, which is also known as "calcifying epithelioma of Malherbe." The most accused etiological factor is repeated skin trauma. Physical examination findings are non-tender, firm, palpable breast mass. The inflammatory phenomena with or without skin ulceration may also be seen. Radiologically, pilomatrixoma is seen as nodular opacity with punctate or heterogeneous calcifications, in mammography and mostly well-defined, circumscribed, parallel-oriented, heterogeneous mass composed of solid-cystic areas and hyperechoic calcifications with acoustic shadowing in US. Internal vascularity in Doppler imaging may also be seen [24, 25] (Fig. 9e, f).

#### Hemangioma

This rare vascular lesion of male breast is formed by proliferation of vascular channels lined by endothelial cells. It is located in either dermal or subcutaneous layers. Hemangioma may be asymptomatic or may present as palpable superficial breast mass, on physical examination. Imaging findings seen in mammography are well-defined, ovoid or lobulated, superficial mass with high density and internal coarse calcifications representing phleboliths. Sonographic findings are well-defined, ovoid or lobulated, superficial, non-hyperechoic lesion with an abrupt interface, with or without internal complex structure. Rarely, these vascular lesions could be seen as hyperechoic mass with indistinct margins [7, 26].

#### Schwannoma

This rare neoplasm arising from Schwann cells are seen as smooth, painless, soft mass, on physical examination. NF-2 related types have higher malignity potential than sporadic types. The mammographic findings are welldefined, round or oval mass. Well-defined, hypoechoic, solid mass with acoustic enhancement are typical sonographic findings. The other sonographic findings are complex heterogeneous mass with central cystic component and internal vascularity [6].

#### Granular cell tumor

Granular cell tumor is a rare benign neoplastic tumor arising from perineural Schwann cells of the peripheral nerves. Physical examination findings are painless, solid, non-tender nodule in the upper inner quadrant of the breast. These nodules can be multifocal. Imaging findings are non-specific and variable with a wide-range spectrum from "circumscribed mass with posterior acoustic enhancement" to "ill-defined mass with posterior acoustic shadowing." They can mimic primary breast cancer [27].

#### Benign non-neoplastic breast lesions Epidermal inclusion cyst

It is the third most common benign lesion of the male breast, which arises from occluded hair follicles or sites of previous skin trauma. Physical examination findings are typically palpable, smooth, superficial lesion, and gland orifice which could be seen as a blackhead over mass. Furthermore, tenderness may be present in cases of rupture and surrounding inflammation. Epidermal inclusion cysts are seen as circumscribed, round or oval, superficial mass adjacent to skin with high density, in mammography. Tangential views can confirm the superficial location of the lesion. Sonographic findings are superficial (within dermis or subcutaneous fat), circumscribed, oval, hypoechoic mass, sometimes with "Onion skin" appearance which represents alternating concentric hypo and hyperechoic rings due to the accumulation of lamellated keratin. Rarely fistula tract can be seen between lesion and skin, in US [3, 15].

#### Sebaceous cyst

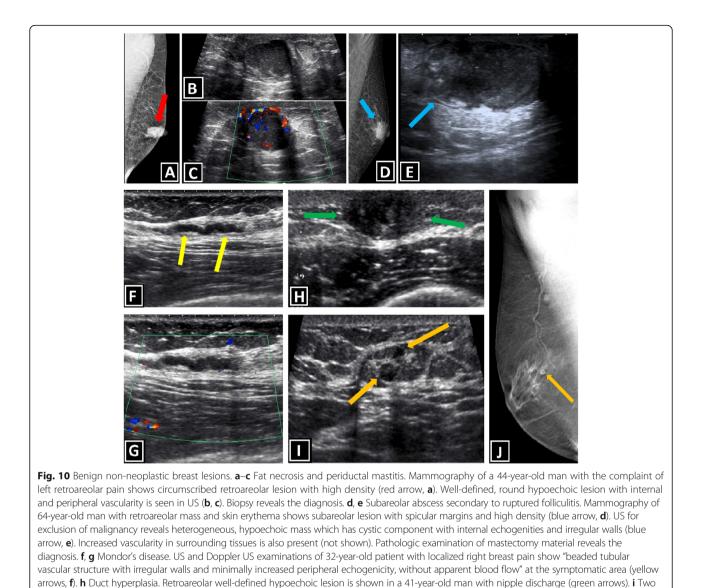
Sebaceous cyst is benign intradermal lesion formed by obstruction of sebaceous gland. It is generally smaller than epidermal inclusion cyst. Sebaceous cyst is one of the "don't touch" lesions when suspected because biopsy may induce inflammatory process in the surrounding breast tissues. Clinically, sebaceous cyst is seen as palpable, superficial breast lump. Inflammatory changes may be present in cases of rupture or surrounding inflammation. The imaging findings are similar to epidermal inclusion cyst, and mostly indistinguishable with imaging alone. The "claw sign," which refers to wrapping of dermal tissue around the lesion like a claw, may be seen in US and could be helpful for differentiation from superficial breast parenchymal lesions. In Doppler US imaging, they do not have internal vascularization with the exception of peripheral hypervascularization which is seen in inflamed or ruptured cases [6, 7, 28].

#### Post-traumatic hematoma and fat necrosis

Fat necrosis is formed by necrotic adipocytes surrounded by lipid-laden macrophages. Hematoma is a localized collection of blood and blood products outside the blood vessels. These are benign processes, generally related to trauma or breast surgery. The clinical presentation is usually new palpable breast lesion which could be painful. The history of trauma, surgery, or anticoagulation is generally present. Mammographic imaging findings are related with the phase of lesion. Acute forms can mimic malignity or abscess due to ill-defined mass with skin thickening, and trabecular prominence. Chronic forms turn into more discrete lesions which may contain fat-fluid levels. Distortions, dystrophic calcifications, and persistent well-defined masses are common mammographic findings during this stage. Sonographic findings are avascular, circumscribed, complex, heterogeneous mass with possible fluid-fluid levels, and internal septation [3, 6] (Fig. 10a–c).

#### Seroma

This lesion usually develops after surgery or breast trauma, and it may be seen as painful mass on physical



small intramammary lymph nodes with circumscribed margin and round shape are shown (orange arrows). j Breast tomosynthesis of another patient

reveals gynecomastia accompanied by intramammarian lymph node with round shape and radiolucent fatty hilus (orange arrow)

examination. An anechoic fluid collection with or without internal septation is the initial imaging finding. During maturation, a collection with thick, nodular margin can be seen [29].

#### Nodular fasciitis

It is one of the benign mesenchymal tumors characterized by pseudoneoplastic reactive proliferation of myofibroblast. The breast involvement is rare. The etiology is unclear while local trauma is accused for triggering reactive proliferation by the majority of authors. Physical examination findings are hard, well-defined, mobile, painless, palpable mass which shows rapid growth with sudden onset, generally, and may mimic malignancy. Radiologically, nodular fasciitis is seen as ill-defined density with irregular margins, without calcifications, in mammography and hypoechoic, heterogeneous, solid mass with irregular margins, without prominent vascularization, in US [6, 30, 31].

#### Diabetic mastopathy

This rare breast disease is characterized by stromal sclerosis, and dense lymphocytic infiltration containing fibroinflammatory process. It is typically seen in patients with long-term type 1 diabetes mellitus. Diabetic mastopathy may mimic malignity both radiologically and clinically. That is why, the history of long-standing diabetes mellitus is crucial for diagnosis. Generally, systemic complications of diabetes coexist with diabetic mastopathy due to long-term disease. There is usually large, painless, firm, palpable breast mass with multicentric or bilateral involvement on physical examination. Radiologically, ill-defined mass or asymmetric densities are usual mammographic findings. Sonographic findings are hypoechoic mass or masses with irregular margins, and posterior acoustic shadowing [6].

#### Periductal chronic inflammation and ductal ectasia

It is a rare disease of the male breast characterized by chronic inflammation, and fibrosis around dilated ducts which may contain debris. It can mimic malignity due to its clinicopathologic properties. Nipple discharge, subareolar tender breast mass, and nipple retraction can be seen on physical examination. Additionally, it could be complicated with subareolar abscess or fistula formation. Common mammographic findings are subareolar ductal dilatation with intraductal calcified secretions, and coexistent mass-like opacities in subareolar location. Retroareolar dilated ducts which contain internal debris is seen as tubular structures in US. So the differentiation from papilloma or papillary carcinoma may be required. Subareolar mass is another sonographic finding which could be present [32, 33] (Fig. 10a–c).

#### Subareolar abscess

It is also called as "Zuska's disease," and it is characterized by subcutaneous abscess in subareolar location due to aseptic inflammation caused by squamous metaplasia which is secondary to obstruction of lactiferous ducts. Physical examination findings are mastalgia, nipple discharge, inflammatory signs, tender subareolar breast lump. Fistulas can be seen in chronic cases. Diagnosis usually requires the knowledge of clinical history and physical examination. Because imaging findings may mimic malignity or gynecomastia. Mammography is hard to obtain due to pain with breast compression. When it is done, imaging findings like skin thickening, distortion, and ill-defined subareolar mass accompanied by trabecular thickening could be seen. Sonographic imaging findings present as "heterogenous, irregular, hypoechoic mass" or "fluid collection with internal echoes, and irregular walls." The increased vascularity in surrounding tissues, thickening of skin due to inflammation, and fistulous tract formation may also be present [6, 34, 35] (Fig. 10d, e).

#### Idiopathic and specific granulomatous mastitis

These are very rare chronic inflammatory male breast diseases characterized by non-caseating granulomatous lobulitis. They may mimic breast cancer both clinically and radiologically. The etiology of idiopathic granulomatous mastitis is not clearly known. While specific granulomatous mastitis usually has certain etiology like tuberculosis, Wegener's granulomatosis, vasculitis, sarcoidosis, foreign body reaction, syphilis, corynebacterial infection, and fungal and parasitic infections. Identification of etiology is important for both prognosis and treatment. Clinico-radiologic findings of these diseases are similar to each other. Physical examination findings are subareolar, firm, painless, or tender breast mass. Inflammatory symptoms may be present, and they are usually resistant to empiric antibiotic therapy without relief. Mammographic imaging findings are ill-defined opacities, parenchymal distortion, and masses with non-distinct margins. Ill-defined hypoechoic heterogeneous lesions which may show tubular extensions, and increased vascularization accompanied by peripheral distortion are known sonographic findings [36–38].

#### Mondor's disease

It is rare self-limiting breast lesion characterized by superficial thrombophlebitis of the anterior chest wall including the superior epigastric, lateral thoracic, thoracoepigastric veins. The exact etiology or the pathogenesis is not known. The most accused factors are direct trauma to the veins, invasive procedures like surgery and biopsy, inflammatory process, and breast cancer. The expected physical examination finding is initially soft and red cord-like structure which later turns into painful and rigid breast mass. Radiologically, Mondor's disease is seen as dilated tubular density beneath the skin in mammography and tubular, beaded hypoechoic or anechoic structure with thick hyperechoic wall in US. There is usually no flow in this tubular structure in Doppler US scan [8, 39] (Fig. 10f, g).

#### Masson's tumor

This rare breast lesion is caused by organization and recanalization of preexisting thrombus with benign vascular proliferation. It is also known as papillary endothelial hyperplasia (PEH). During the organization of thrombus, anastomosing vascular channels develop and lesion becomes prominently vascular. Because of this, angiosarcoma, which is the most frequent vascular malign tumor of the breast, is the important differential diagnosis and it must be excluded. However, differentiation of Masson's tumor from angiosarcoma is not possible always, even pathologically. The imaging findings are nonspecific [40].

#### Ductal hyperplasia

Ductal hyperplasia is defined as intraductal epithelial cell proliferation with or without atypia. It could be seen incidentally in gynecomastia. Although, physical examination findings are non-specific, nipple discharge or palpable mass may be seen. Imaging findings of ductal hyperplasia have been rarely described. The cluster of amorphous microcalcification may be seen in mammography, especially in case of atypical ductal hyperplasia. Well-defined nodular lesion or area of distortion may be present in US [41-43] (Fig. 10h).

#### Venous malformation

It is the most common type of vascular malformation in the male breast, consisting of multiple ectatic thinwalled vessels lined by endothelial cells. The slow internal flow without shunting is a characteristic feature. Common physical examination findings are unilateral, painless, chronic breast enlargement. Venous malformation is seen as multiple tubular densities with or without phleboliths in mammography, while multiple tubular, anechoic spaces with internal vascularity in US. The slow venous flow in Doppler US is a very important finding for differential diagnosis, since lack of flow suggests the lymphatic malformation. The phleboliths and thrombosis are other findings that can be detected sonographically [6].

#### Intramammary lymph node (IMLN)

IMLNs can be anywhere but generally located in upperouter quadrant, and the most common physical examination findings are palpable, mobile breast mass. Imaging findings of the IMLN are "oval or reniform lesion with radiolucent fatty hilus and denser peripheral cortex" in mammography and "echogenic hilus with thin (<2-3mm), homogenous, hypoechoic cortex" in US. Sonographically, short-axis dimension > 10 mm, round or irregular shape and cortical thickening (>2-3 mm) is considered as abnormal [6] (Fig. 10i, j).

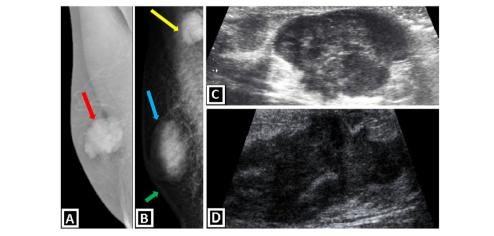
#### Breast augmentation

Augmented breasts have diffuse gynecomastia and female breast appearance due to hormonal therapy. "Surgically placed breast implants" and "direct injection of viscous fluids: mineral oils like industrial silicone or paraffin" are other alternatives, in transexual patients. The mineral-oil injection may lead to inflammation and necrosis of breast. Long-term complication of the mineral-oil injection is known as "sclerosing lipogranulomatosis." Physical examination findings are inflammation, fibrosis, and palpable mass-like lesions (sclerosing lipogranulomas). Mammographic findings are various such as "Female breast" appearance with extensive microcalcifications and/or coarse calcifications, parenchymal opacities and distortions. Diffuse gynecomastia, breast implants, calcifications with acoustic shadowing, distortions due to fibrosis or inflammatory breast masses can be seen in US imaging [6, 44].

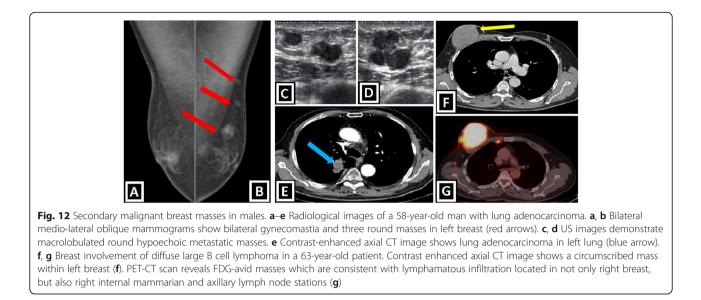
#### Malign breast lesions

#### Invasive ductal carcinoma (IDC)

It is the most common primary malign neoplastic lesion of male breast, and almost 80% of all cases belong to IDC "not otherwise specified type." On physical examination, these lesions are seen as hard, painless, palpable mass with secondary features such as nipple retraction, skin thickening, palpable axillary LAP. The latter one accompanies IDC in 50% of the cases. The discrete mass appearance and secondary features are important clues for malignity. Bilateral mammography is recommended in case of malignity suspicion, because underlying risk factors may cause the development of contralateral breast malignity. In mammography, IDC is seen as radiodense, irregular, retroareolar masses with spiculated, lobulated or microlobulated margins. The incidence of microcalcifications is lower than female breast cancer. Retroareolar, non-parallel, hypoechoic mass with irregular borders and variable vascularity are well-known



**Fig. 11** Mammography and US images show primary male breast cancer. **a**, **b** Medio-lateral oblique mammograms show invasive ductal carcinoma of the breast. Lobulated subareolar masses with high density, without microcalcification are seen (red and blue arrows). **a** Grade 2 invasive ductal carcinoma. **b** Grade 3 invasive ductal carcinoma with axillary lymph node metastasis (yellow arrow) and skin thickening (green arrow). US images of grade 2 (**c**) and grade 3 (**d**) invasive ductal carcinomas reveal irregular shape and indistinct margins of the masses



US findings. Accompanying secondary findings can also be seen in US [3, 4, 7, 9, 15] (Fig. 11).

#### Ductal carcinoma in situ (DCIS)

It is the second most common subtype of male breast cancer and responsible for 5% of all cases. The pure form of DCIS is rarely present due to the very high rate of coexistence with invasive carcinoma. Palpable mass and/or bloody nipple discharge is frequently seen in physical examination. DCIS is seen as pleomorphic microcalcifications which represent in situ component of tumor cells, in mammography. However, parenchymal opacity or distortion may, also, be seen if invasive carcinoma accompanies. The findings on US are non-specific, but coexistent IDC foci may be seen as mass or distortion [6, 7, 9].

#### Papillary carcinoma

Papillary carcinoma is more common in men than women (2:1) accounting for 2.5–5% of male breast cancers and presents with palpable subareolar mass. Mammographic findings are circumscribed, oval, lobulated or irregular, subareolar mass. Ill-defined borders point out infiltrative component. Typical sonographic appearance is complex heterogeneous mass formed by solid and cystic components. "Solid mural nodule in complex mass" or "pure solid mass" are other possible findings [3, 7].

#### Invasive mucinous carcinoma

IMC is a histological variant of breast carcinoma characterized by the extracellular mucin component surrounding neoplastic cells. It accounts for 1% of male breast cancers. Pure and mixed forms are present. Physical examination findings are mostly non-specific. Palpable hard subareolar mass may be seen. Imaging findings are, also, non-specific. Pure form may be confused with benign breast lesions, because the characteristic mammographic finding of pure form is round and welldefined opacity, and the expected sonographic finding is well-defined subareolar iso-hypoechoic lesion [7].

#### Dermatofibrosarcoma protuberans

This uncommon lesion, which is locally aggressive, slowgrowing fibrous tumor involving the dermal layer of the skin with intermediate-to-low grade malignancy, is more frequent in males than females. Due to its local aggressive characteristics, local recurrence rate after the surgery is very high. It presents as firm, painless, slow-growing, generally mobile, palpable breast lump, on physical examination. Mammographic findings are ill-defined radiodense breast mass without fat or calcification. An ovoid lesion with lobulated or microlobulated margins located in dermis or subcutaneous tissue is sonographic appearance. Mixed internal echogenicity and hypervascularization can also be seen in US [15, 45, 46].

#### Paget's disease

This disease is characterized by skin changes of nipple and areola, such as eczematous scaling, skin erosions, and ulceration associated with the possible ductal spread of breast cancer. It is mostly a clinical diagnosis, and imaging findings are non-diagnostic. However, coexistent DCIS and/or invasive carcinoma may be present, and their imaging findings can be seen. After suspicion "based on history and physical examination," biopsy confirms the diagnosis [7].

#### Lymphoma

Lymphoma involvement of male breast may be primary or secondary. Secondary cases mostly related to non-Hodgkin B cell lymphoma involvement. Physical examination findings are enlarged axillary lymph nodes, single or multiple palpable breast masses. The history of lymphoma is an important clue for the diagnosis. Mammographic imaging findings are "single or multiple, circumscribed or ill-defined lesions" and "multiple, enlarged, circumscribed, oval or lobular, radiodense axillary lymph nodes without apparent radiolucent fatty hilus". Findings in US imaging are "circumscribed or irregular, hypoechoic, solid mass" or "masses accompanied by enlarged axillary lymph nodes with irregularly thick-ened cortex and without normal echogenic fatty hilus structure" [1, 3, 7, 15] (Fig. 12f, g).

#### Metastases

Breast metastases from extramammarian malignancies are rare, and they point out late-stage disseminated disease. The presence of multiple bilateral lesions and subcutaneous fat tissue location are supportive features for metastases, while "unilateral lesion" and "glandular breast tissue involvement" support primary breast cancer. Regarding breast metastases, skin or nipple retraction is not expected in contrast to primary breast cancer. Multiple bilateral palpable solid masses with a known primary malignancy (such as prostate cancer, lung cancer, gastric cancer, colorectal cancer, melanoma, or sarcoma) suggest the possibility of breast metastases. Prostate cancer is accepted as the most frequent primary site for breast metastasis. Mammographic imaging findings are single or multiple (mostly), round or lobulated masses with circumscribed or ill-defined (less expected) borders. The diffuse skin thickening may, also, be seen. Calcifications and spiculated borders are not expected findings opposite to primary breast cancer. Main imaging findings in US are hypoechoic, circumscribed masses. Increased internal vascularity and posterior acoustic enhancement are other features that may be present [1, 4, 18] (Fig. 12a-e).

#### Conclusion

Understanding the histoanatomic differences between male and female breasts has utmost importance for insighting the diversities in radiological appearance, diagnostic algorithm, biopsy procedure, and malignancy characteristics, between the two sexes. Majority of breast lumps in male patients are benign, and the most common reason is gynecomastia. Except for gynecomastia, male breast has wide-range spectrum of pathologies and hosts many lesions which can be classified according to various radiologic and histopathologic characteristics. Familiarity with the imaging findings, histopathological properties, and presentation features of various benign and malignant male breast lesions allows correct imaging interpretation which can facilitate "convenient patient management," "reaching early and confident diagnosis," and "avoiding unnecessary interventions".

#### Abbreviations

DCIS: Ductal carcinoma in situ; FDG: Fluorodeoxyglucose; IDC: Invasive ductal carcinoma; ILC: Invasive lobular carcinoma; IMLN: Intramammary lymph node; LAP: Lymphadenopathy; LCIS: Lobular carcinoma in situ; MRI: Magnetic resonance imaging; NF: Neurofibromatosis; PASH: Pseudoangiomatous stromal hyperplasia; PEH: Papillary endothelial hyperplasia; PET-CT: Positron emission tomography-computed tomography; TDLU: Terminal ductal-lobular unit; US: Ultrasound

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Not applicable

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#### **Competing interests**

The authors declare that they have no competing interests.

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# **Male Breast Disease**

- 1. According to the text, the majority of the male breast lumps are benign, and the MOST common reason is \_\_\_\_\_\_.
  - A. injury
  - B. an increase in testosterone
  - C. gynecomastia
  - D. obesity
- 2. The male breast is located in between the 2<sup>nd</sup> and \_\_\_\_\_ ribs.
  - A.  $6^{\text{th}}$
  - B.  $7^{th}$
  - C. 8<sup>th</sup>
  - D. 9<sup>th</sup>
- 3. True or false. The mammary glands of both sexes are identical at birth.
  - A. True
  - B. False
- 4. Involution and ductal atrophy occur in the male breast due to significant increase
  - in \_\_\_\_\_ levels.
    - A. estrogen
    - B. testosterone
    - C. progesterone
    - D. prolactin
- 5. According to the text, at the time of male breast cancer diagnosis, axillary lymphadenopathy (LAP) involvement is at nearly \_\_\_\_\_ percent.
  - A. 30
  - B. 40
  - C. 50
  - D. 60
- 6. According to the text, mammography is the first-line suggested imaging modality in case of suspicious physical examination of male breast lesion, with \_\_\_\_\_\_ as the next recommended step when mammography findings are inconclusive.
  - A. magnetic resonance imaging (MRI)
  - B. computed tomography (CT)
  - C. nuclear medicine
  - D. ultrasound (US)
- 7. The main anatomic difference between diffuse gynecomastia and female breast is the absence of \_\_\_\_\_\_ ligaments.
  - A. Conner
  - B. Cooper
  - C. Braxton
  - D. Alexander
- 8. Pseudogynecomastia is mostly seen in overweight-obese individual or with those with neurofibromatosis type-\_\_\_\_\_ history.
  - A. 1
  - B. 2
  - C. 3
  - D. 4

- 9. Gynecomastia has bi-modal age distribution involving the peripubertal period and the period after age \_\_\_\_\_ years.
  - A. 35
  - B. 40
  - C. 45
  - D. 50
- 10. According to histopathological type and behavior, male breast lesions can be divided into \_\_\_\_\_\_ main categories.
  - A. two
  - B. three
  - C. four
  - D. five
- 11. According to the text, \_\_\_\_\_\_ is the second **MOST** common benign male breast lesion.
  - A. cyst
  - B. hematoma
  - C. lipoma
  - D. papilloma
  - \_\_\_\_\_ is a slow-growing benign neoplastic lesion arising from brown
  - adipose tissue.

12.

- A. Pilomatricoma
- B. Hibernoma
- C. Intraductal papilloma
- D. Schwannoma
- 13. According to the text, epidermal inclusion cysts are the \_\_\_\_\_ MOST common benign lesion of the male breast.
  - A. second
  - B. third
  - C. fourth
  - D. fifth
- 14. Subareolar abscess is also known as "\_\_\_\_\_" disease.
  - A. Zuska's
  - B. Manor's
  - C. Mondor's
  - D. Riding's
- 15. Ductal carcinoma in situ (DCIS) is responsible for \_\_\_\_\_ percent of all male breast cancer cases.
  - A. 5
  - B. 10
  - C. 15
  - D. 20

	X-RAY CE	
Male Breast Disease 1.5 A CEU	J's Answer Sheet	
First Name	Last Name	
State License # (If applicable)	ARRT #	
Address		
Phone		
Would you like your certificate	e faxed or emailed back to you? Yes / No	
Fax number		
Email		
Please darken th	e answers you feel are correct.	
<b>1.</b> (A) (B) (C) (D)	-	

(A) (B) (C) (D)
(A) (B) (C) (D)