



Using Breast MRI to Evaluate Complications of Autologous Fat Grafting

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Abstract

Introduction: During the past few years in Taiwan, there has been a rapid growth of autologous fat grafting procedures performed for cosmetic reasons. We retrospectively demonstrate Complications of Autologous Fat Grafting in the breast, based on clinical presentations and breast MRI imaging. We also discuss the possible method of early detection of complications.

Material and method: Twenty-one ethnic Chinese (Taiwanese) women were transferred for delicate breast MRI study in the past year. These patients had suffered different signs and symptoms following the procedure. Clinical and image information were collected from the hospital database.

Results: Breast MRI successfully detected all viable fat tissues and complicated lesions of the breasts. The image findings corresponded to the clinical presentation and were used in therapeutic decision making. Fat necrosis with granuloma formations caused by graft rejection was the main complications. The most severe complication was an abscess formation and impending sepsis. Management procedures included operation or drainage; received oral antibiotics; and received clinical observation.

Discussion and Conclusion: MRI is capable of detecting the surviving fat graft tissues, and to distinguish degree of malignancy from benign lesions. In complicated cases, the highest incidences showed granuloma formations and fat necrosis. However, laboratory data such as CRP, IgE and complement (C3 &C4) levels only reflected some of the infectious cases and could not be used as monitoring parameters. Breast MRI is an ideal tool for early detection of autologous fat complications without causing indistinguishable conclusions from malignancy. Breast MRI should be performed as early as possible for this complication, so that help for early treatment can be planned.

Keywords: Breast MRI; Autologous fat graft; Survive; Abscess formation; Fat necrosis; Rejection; Complication

Introduction

Autologous fat grafting is a technique that involves using the patient's own adipose tissue to be applied on a subcutaneous area of the body to increase total fat volume [1-3]. Breast augmentation using autologous fat grafting is not associated with implant related problems such as implant leakage, breast deflation, visible or palpable implants, or the development of breast capsular contracture. Regarding successful cases of autologous fat grafting, under the condition of suspected tumor growth, a usual biopsy procedure can be performed without risk of a stabbing rupture of the implanted materials. However, autologous fat grafting in the breast is not a simple procedure and should be performed by highly trained and skilled surgeons. The recipient of autologous fat grafting is at risk of intraoperative pneumothorax, calcification, multiple cyst formation [4], focal breast indurations, pain, infection, and abscesses [5], abnormal breast discharge, and reactive lymphadenopathy. In contrast to the aforementioned successful cases, these patients require life-long follow up for breast cancer screening because of the difficulty of cancer detection. These patients should also undergo long-term imaging analyses [6,7]. Autologous fat grafts are also believed to have the disadvantage of the unpredictability graft tissue survival because of fat necrosis and re-absorption [8]. The incidence of calcifications (mostly due to fat necrosis) after all types of breast operations varies, but has been reported in as many as 50 percent of patients within 2 years [9].

It was also reported that breast asymmetrical densities were present in approximately half of patients throughout the follow-up period of post-mammoplasty. In additions, fat necrosis occurred in approximately one tenth of the patients [9]. Many plastic surgeons have rejected fat grafting in the breast because of the speculation that transplanted fat might undergo necrosis or die off gradually, forming lumps or calcifications. Fat grafting raises the concern of interfering with breast cancer detection. In fact, injecting small volumes of fat produces microcalcifications, which can be indistinguishable from carcinomas in mammograms; whereas injecting larger volumes might result in the liponecrotic pseudocyst formation that can interfere with physical breast examination and cause a growing and painful mass [10,11]. Regarding breast image screening and diagnosis for such patients, mammography is favored by radiologists to differentiate cancer microcalcifications from benign calcifications of the breast. However, most cosmetic mammoplasty patients are among the young age group that should receive non-ionized radiation screening of breast tissues. Therefore, the first line screening for this age group is usually sonography, but the sensitivity and specificity of sonography to differentiate calcification and microcalcifications is not reliable [12]. Breast MRI should be considered a first line screening tool or

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at least used when clinical suspicion or radiographic abnormality is indeterminate.

Breast MRI has also been reported to be more capable than mammography of early detection of fat necrosis and clinically nondetectable masses and nodules, although MRI is not used to detect calcifications or microcalcifications [13].

During the past 2 years in Taiwan, there has been a rapid growth of autologous fat grafting procedures performed for cosmetic reasons. In this study, we demonstrate cases of autologous fat grafting in the breast, including complications, based on clinical presentations and breast MRI imaging. We also discuss the possible method of early detection of the Complications of Autologous Fat Grafting of the breast.

Material and Methods

Cases collections

From March 2011 to February 2012, twenty-one ethnic Chinese (Taiwanese) women were transferred to the Breast Health Center (Taipei Medical University Hospital) for delicate breast MRI study. These patients exhibited different symptoms and signs after the procedure. Clinical and image information were collected between March 2011and February 2012 from the hospital database. The patients were between 25 and 62 years old, and none had undergone treatment in the period between mammoplasty and breast MRI study. Furthermore,

information collected and recorded included clinical expression, laboratory data such as Immunoglobulin E (IgE), Antinuclear Antibodies (ANA), complement (C3&C4), C-Reactive Protein (CRP), antinuclear antibodies (ANA), monocyte and bacteria culture, as well as the possible pathological diagnosis and method of treatment.

Image processing using delicate breast MRI

Pre-enhanced and post-enhanced subtracted images were obtained, and an enhanced curve analysis was performed. All imaging studies were performed on a 1.5 T dedicated spiral breast MRI system (Aurora Systems, USA) [14-17]. The breasts were fitted and positioned using a breast transmission-receiving (TR) coil. The off-resonance spiral image mode was used. A gradient echo with 10 ms of TR and T2-weighted spin echo with 2.5 ms of TR were acquired. The echo time was 5 ms for RODEO and 120 ms for the T2 sequences. The volumetric matrix size was 360x360x128 mm. The slice thickness was 1.1 mm. The field of view of each station was 20–36 cm. Sequences were performed before and after the infusion of 0.2 mmol/kg adolinium-diethylenetriamine pentaacetic acid (Magnevist; Bayer-Schering Pharma AG, Berlin, Germany), administered as a bolus dose with a power injector, followed by a 20-mL saline flush. Subtraction images were also performed [14-17].

Results

According to our data on these 21 patients (Table 1), the complications may have developed 12 days to 24 months later after

Case No.	Age	Period of autologous fat graft	Symptoms and signs	Image findings on MRI	Visibility of grafted fat	Laboratory data	Clinical or pathological diagnosis	Treatment
Case 1 (Figure 1)		`12 days	Pain and fever	(1)Right breast enhanced cavities (2)Ipsilateral lymphadenopathy (>2cm)	Yes	(1) Anemia	(1) Abscess	Operation and drainage
	35					(2) Significant CRP elevation	(2) Impending sepsis	
						(3) Bacterial culture:Candida albicans isolated		
Case 2 (Figure 2)	30	4 months	Right breast tenderness with lump	(1)Multiple granulomas and abscess formation at deep superior and subareolar region of right breast. The largest one is over 3.2cm.	Yes	(1) IgE elevation	Mastitis and granulomatous formation	Operation and drainage
						(2) Monocyte increase		
				(2)Mild inflammation of pectoral major muscle.		(3) CRP normal		
						(4) Bacterial Culture: No growth		
Case 3 (Figure 3)	27	3 months	Tenderness at bilateral sides	(1) Diffuse granuloma formations, pockets of thick wall pockets with various sizes, within glandular tissues of bilateral breasts.Suggest process of fat necrosis and non-infectious inflammatory process	Yes	Nil	Mastitis and granulomatous formation	Observation
				(2)Nonspecific enlarged LNs at bilateral breast axillary regions.				
Case 4 (Figure 4)	51	24 months	(1) Tenderness at bilateral sides	(1)Multiple small ill defined encapsulate fat masses, most of the masses exhibit wall enhancement.	Yes	Nil	Mastitis and granulomatous formation	Oral Antibiotics
			(2) Middle chest pain	(2)Focal inflammatory lesion at anterior of sternum, with 1.8cm.				
				(3) Non specific reactive axillary LNs.				
Case 5 (Figure 5)	25	8 months	Palpable lumps at superior of right breast and medial of left breast	1) Diffuse granuloma formations , pockets of thick wall pockets with various	-Yes	Nil	Mastitis and granulomatous formation	Observation
				sizes, within glandular tissues of bilateral breasts.				
				Suggest process of fat necrosis and non- infectious inflammatory process				
				(2)Nonspecific enlarged LNs at bilateral breast axillary region.				

Page 3 of 12

Case 6 (Figure 6)	44	1 months	Palpable lumps at superior of right breast	(1)Multiple granulomas like masses at right breast. Suggest ongoing process of inflammation with fat necrosis,	Yes	Nil	Mastitis and granulomatous formation	Oral Antibiotics
				(2) Possibility of mild infection is also considered.				
Case 7 (Figure 7)	26	12 months	Palpable lumps	(1)Multiple granulomas like masses at bilateral breasts.	Yes	(1) IgE elevation	Mastitis and granulomatous formation	Observation
				(2)Obvious abnormal LNs(>1.5cm, dense) at bilateral breasts axillary regions. (1) Right breast mass; and other smaller		(2) Antinuclear Antibodies (ANA): 1:40 (elevation)		
						(3) Significant (complement,C4) :elevation		
						(4) Normal CRP		
Case 8 (Figure 8)	62	12 months	Palpable lumps at right breast	nodular lesions in bilateral breasts	Yes	Nil	Mastitis and granulomatous	Operation
				(2) Probably chronic inflammation due to fat grafting induced granuloma formations			formation	
	36		Inferior lump	(1)granulomas and	Yes	Nil		
Case 9 (Figure 9)		6 months	at right; Bilateral medial pain for 2 weeks	fat necrosis. (2)Suspect status of inflammatory infectious			Mastitis and granulomatous formation	Oral Antibiotics
(Figure 9)				or noninfectious				
				(1)Fat grafted cavities at bilateral sides of subglandular regions. (2)Fluid content at left side fat grafted cavities. (3)Some pockets with fluid at inferior of left breast. (4)Post GD	Yes	(1) white cell count:	Mastitis ; acute stage fat necrosis with granuloma formations	
Case 10 (Figure 10)						(2) CRP: normal		Operation and aspiration
	35	6 months				(3) IgE: normal		
				enhanced glandular tissues at left anterior		(4)		
				and bilateral of breast.		Complements(C3&C4): normal		
Case 11 (Figure 11)	35	9 months	Bilateral breast pain after 2 months of fat grafts	 (1)Fat grafted pockets in bilateral sides(one at Rt side and several at left side) of subglandular regions. (2)Minimal fluid content within left side fat grafted pocket. 	Yes	(1) White cell count: normal, but neutrophil predominate(>75%)	Fat necrosis Infectious mastitis	Oral antibiotics
						(2) CRP: elevation		
						(3) IgE: normal		
				3)Enhanced area at left side subareolar region		(4) omplements(C3&C4): normal		
Case 12	37	7 months	Palpable lumps without pain	(1)Post fat grafted and fat necrosis, with cavities within inferior of right breast.	Yes	(1)White cell count: normal, but neutrophil predominate (>75%)	Fat necrosis Non-infectious mastitis	Observation
(Figure 12)						(2)CRP: normal		
				(2) Enhanced area at left side		(3)IgE: normal		
						(4)Complements(C3&C4): normal		
Case 13 (Figure 13)	54	18 months	Palpable lumps without pain	Post fat grafted and fat necrosis, with cavities(like honeycomb) at subglandular of right breast.	Yes	Nil	Fat necrosis	Observatior
Case 14 (Figure 14)	31	10 months	Palpable lumps without pain, at right breast	Enhanced area at right side and reactive lymphadenoapthy	Yes	(1)White cell count: normal	Non-infectious mastitis	Observation
						(2)CRP: normal		
(Figure 14)						(3)IgE: slightly elevation		
						(4)Complements(C3&C4): normal		
Case 15 (Figure 15)	39	13 months	Palpable lumps without pain, at right breast	Enhanced area at right side and reactive lymphadenoapthy	Yes	(1)White cell count: normal		
						(2)CRP: normal	Non-infectious	Observation
						(3)IgE: normal	mastitis :	Observation
						(4)Complements(C3&C4): normal		
Case 16 (Figure 16)	56	5 months	Palpable lumps without pain, at bilateral breasts	Fat necrosis, with cavities within bilateral breasts. Enhanced lesion at deep of left breast	Yes	(1)White cell count: normal	Non-infectious mastitis	Observation
						(2)CRP: normal		
						(3)lgE: remarkable elevation		
						(4)Complements(C3&C4): normal		

Case 17 (Figure 17)	53	4 months	Palpable lumps without pain, at bilateral breasts	There are multiple fat grafted cavities at bil sides of subglandular Regions, with rim enhancements and fluid like signal intensity.	Yes	(1)White cell count: normal	Non-infectious mastitis	
						(2)CRP: normal		
						(3)IgE: remarkable elevation		Observation
						(4)Complements(C3&C4): normal		
Case 18 (Figure 18)	24	11 months	Palpable lump at peri-areolar of Lt breast	Focal enhanced non-mass lesion with irregular border, at subcutaneous skin at anterior lateral inferior of Lt breast. No extensive fat necrosis.	Yes	(1)White cell count: normal	Non-infectious mastitis	Observation
						(2)CRP: normal		
						(3)IgE: normal		
						(4)Complements(C3): slightly elevation		
Case 19 (Figure 19)	30	7 months	Palpable lumps without pain, at bilateral breasts	Multiple fat grafted cavities at superior lateral of bilateral breasts.	Yes	Nil	Fat necrosis with granuloma formations	Observation
			Palpable			(1) Anaerobic Culture	Mastitis, minimal abscess	Oral Antibiotics
Case 20 (Figure 20)	31	3 months	lumps with pain for 2 months, at bilateral breasts	Areas of enhancement with impending abscess formations at superior lateral of right breast	Yes	(2) Aerobic Culture		
Case 21 (Figure 21)	45	6 months	Pain for 2 months and no lump	Reactive lymphadenitis at bilateral axillary regions	Yes	Nil	Non-infectious mastitis	Observation

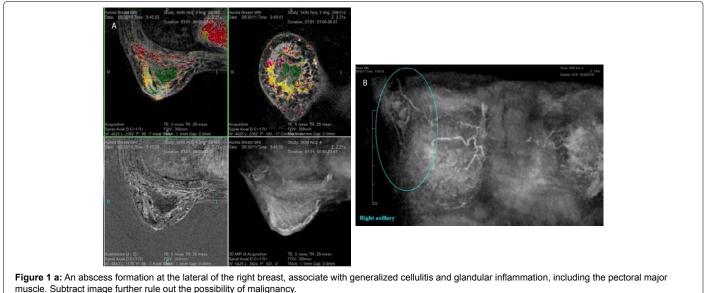
Table 1: Clinical presentation, image finding and treatment of the patients.

autologous fat grafting, including infection, granuloma, and fibrotic change [18-20]. The mean of duration that represent for MRI image diagnosis was 8.1 months. All of the patients suffered from breast tenderness, palpable lumps, pain or fever. The most severe complication was immediate post-operative abscess formation and impending sepsis. All of the cases were diagnosed with fat necrosis and granuloma formations caused by fail of graft tissue survival or graft rejection. Serum test for two of the patients were exhibited CRP elevation, and serum test for IgE was also found elevated in another 3 patients. Among these 21 cases, 19% required operation or incisional drainage; 23.8% received oral antibiotics for conservative treatment; and 57.1% of the cases were suggested for observation after image diagnosis. Based on the MRI images (Figures 1-21), all of the fat grafted tissue from the 21 cases could be identified, even Case 4, who received the procedure for

over 2 years. According to the pictures of breast MRI, no case revealed a lesion indistinguishable from malignancy that required additional biopsy.

Discussion

There are two issues to discuss. Firstly, is that complications of the fat graft can be detected and survived; secondly is breast MRI is capable to distinguish malignancy from benign lesions. In early 2007, Kaufman MR et al reported no evidence of the post autologous fat grafting fat survivability and the predictability of volume restoration. In conducting a quantitative study using large-scale clinical assessment and three-dimensional volumetric imaging to provide useful outcome data, they found that fat survival of autologous fat graft in that period is questionable [21]. By contrast, a rapid growth of new procedures



b: Multiple reactive >2.0 cm axillary lymph nodes.

Page 5 of 12

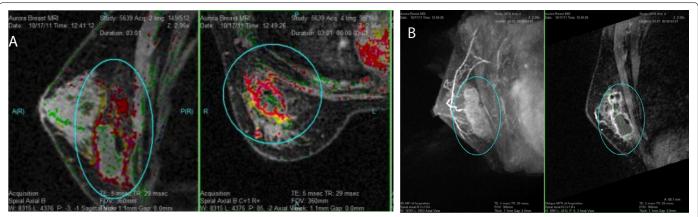


Figure 2: Multiple granulomas and abscess formation at deep superior and subareolar region of the right breast. The largest is over 3.2 cm, with a mild inflammation of the pectoral major muscle.

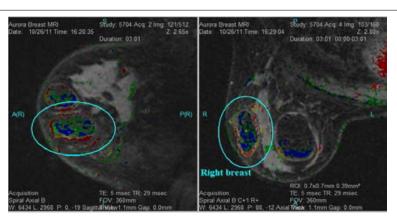
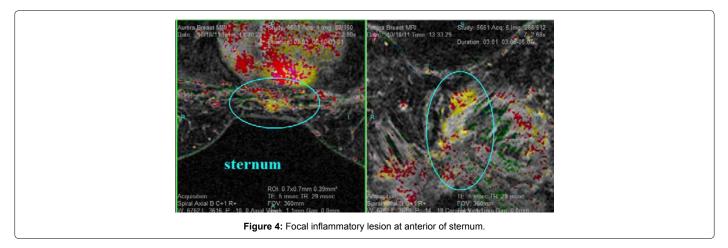
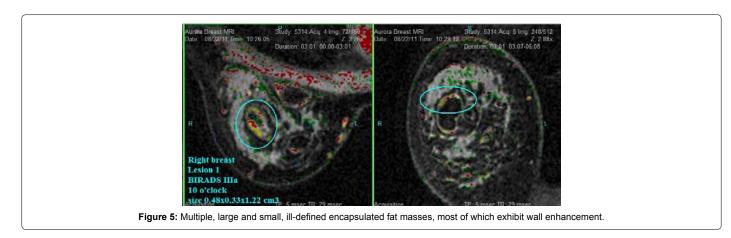


Figure 3: Thick wall pockets of various sizes within glandular tissues of the left breast. On color mapping, there are showed in different color areas represents minimal fluid, fat, protein components.



used to collect larger amounts of transplantable fatty tissue by using liposuction (e.g., water-assisted liposuction); and the procedure of autologous lipo-transplantation after tissue separations. By conducting a series of procedures that focus on the superfluous water, fat purification, and centrifugation, the viable fat tissue could then be harvested [22,23]. Smith et al. using breast MRI to compare the image morphological change of fat grafted breasts before and after using this type of improved technique in operations [22]. They found that 41.1% of the studied patients exhibited a significant volume increase of grafted fat within the treated areas. The average size of increase was approximately 100-150 ml. Approximately 2.4% patients developed palpable subcutaneous nodules which finally proved to be granulomas [22]. For these cases that developed complications, the pathologic results for the excised granulomas, included fat necrosis, calcification, hyalinization, and fibroplasia [24]. Our observations support the results of these previous studies, in that the complication rate of granuloma formation (a complex of chronic inflammation with fat necrosis), is ahead of infection and mastitis. Therefore, granuloma formations due

Page 6 of 12



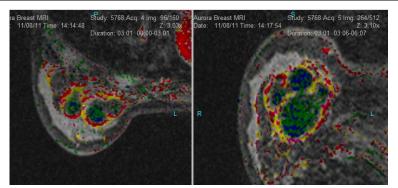
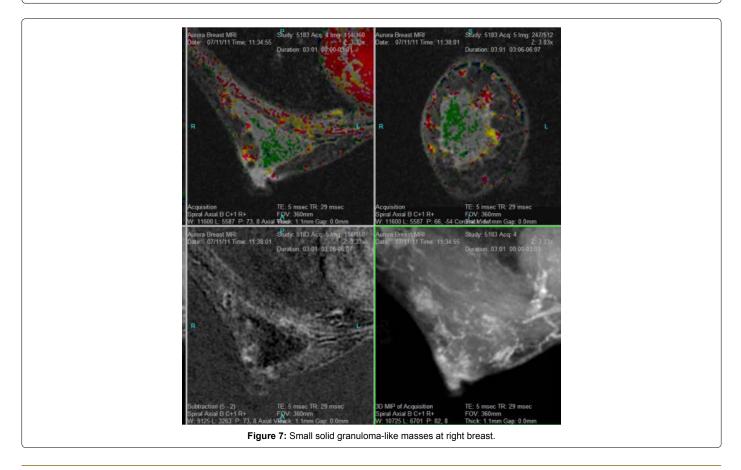


Figure 6: Multiple granuloma-like masses at the subglandular of the right breast. The largest mass (3.0 cm) exhibits wall enhancement with semifluid and semi-fat content.



chronic inflammation of fat tissues, acted as the leading complication type in autologous graft breasts.

Granulomatous formation is caused by the failed attempts of the human immune system to eliminate foreign substances. Granulomas are seen in various diseases that relate to both infectious and noninfectious [25-27]. Granulomatous formation is a type of inflammation, with an important feature that it may contain necrosis. The identification of necrosis in granulomas is important because granulomas with necrosis tend to have infectious causes. However, in most of our cases with granulomatous formations that were associated with fat necrosis, infection did not occur. Fat necrosis has been reported in many different surgical procedures, such as reduction mammoplasty, mastectomy and post reconstructive procedures and others [28-35]. It was also shown that some over-obese animals are predisposed to fat necrosis without trauma and surgical inductions [36,37].

The mechanism of post-operative fat necrosis is still poorly understood but a variety of factors are thought to be some of the causes, such as direct trauma, infection, avascularity, hypoperfusion

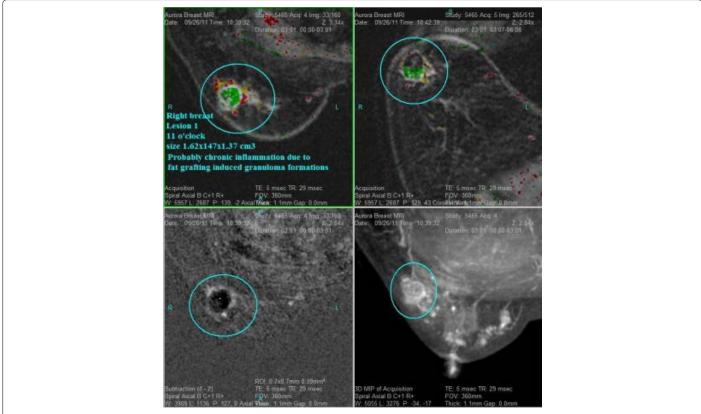
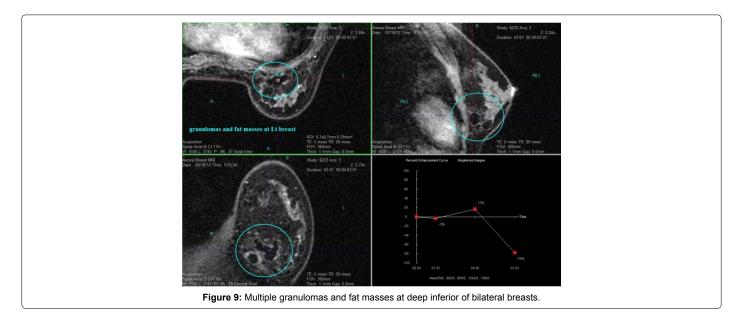


Figure 8: Ill-defined heterogenous enhanced masses at the superior lateral of the right breast ,with wall enhancement.



and lipolytic enzymatic digestion [36-39]. To compare the image features of granulomas under mammography, sonography and breast MRI, it was found the typical mammographic pictures of granulomas as asymmetrical focal densities with no distinct margins, ill-defined masses with spiculated contours, and bilateral multiple ill-defined nodules were seen [40]. Under sonography, granulomas may appear

as nonspecific discrete or heterogenous hypoechoic masses [38-40]. Thus, the mammography and sonography may not provide a conclusive result for determination of the necessity of surgical treatment, or to rule out malignancy [38]. It was believed that a major part of autologous fat injection for breast augmentation would not survive due to the associated physiological response of fat to succumb to necrosis and

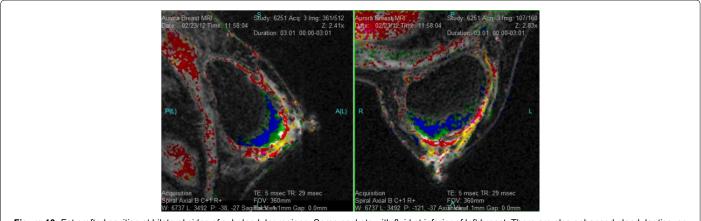


Figure 10: Fat grafted cavities at bilateral sides of subglandular regions. Some pockets with fluid at inferior of left breast. There are also enhanced glandular tissues at left anterior and bilateral lateral of breasts, represent mastitis.

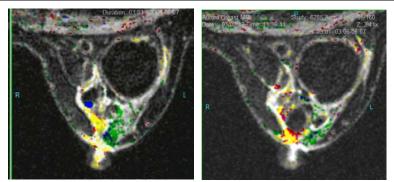


Figure 11: Multiple thick wall enhanced pockets with minimal fluid at inferior of left breast. There are also enhanced glandular tissues at left anterior and bilateral lateral of breasts, represent mastitis.

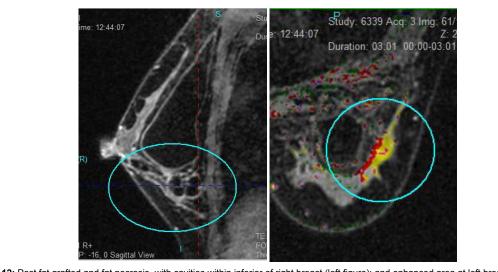
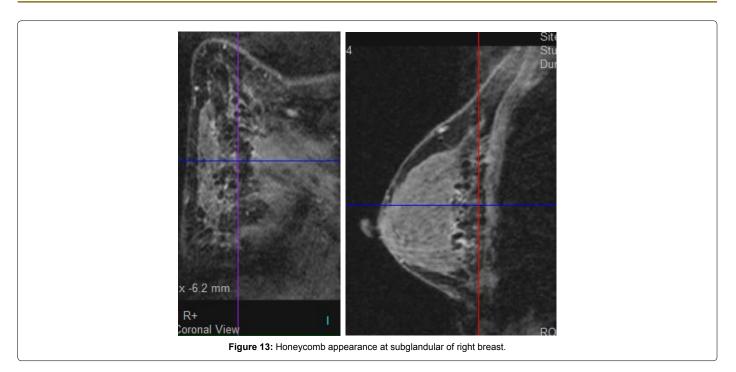


Figure 12: Post fat grafted and fat necrosis, with cavities within inferior of right breast (left figure); and enhanced area at left breast (right figure).

Page 9 of 12



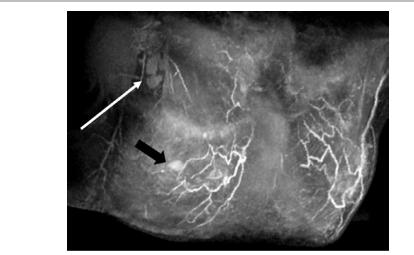
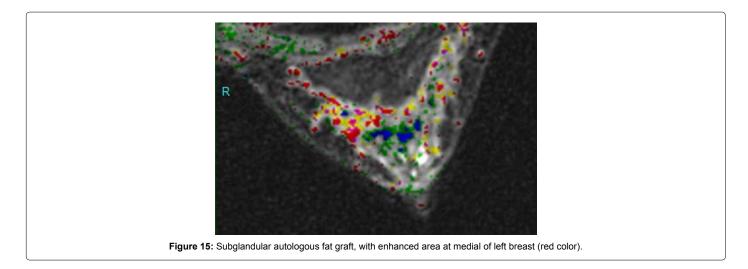
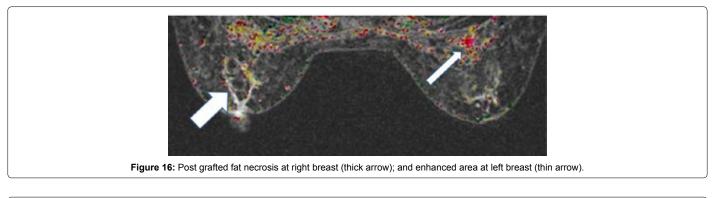
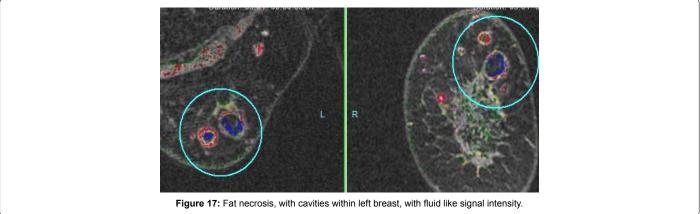


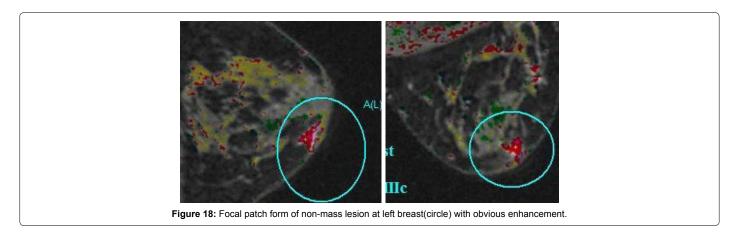
Figure 14: Post GD enhanced mastitis area (black thick arrow) at right breast with and reactive lymphadenopathy (thin white arrow).

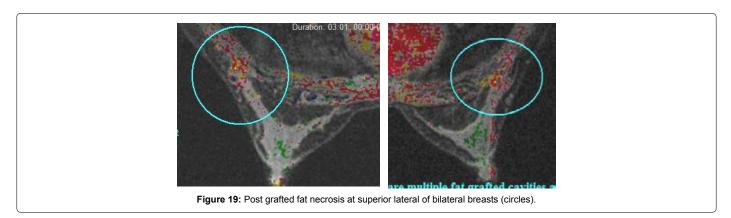


Page 10 of 12









Page 11 of 12

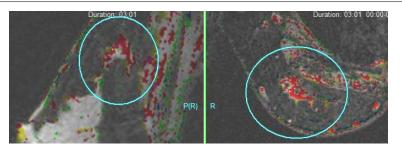
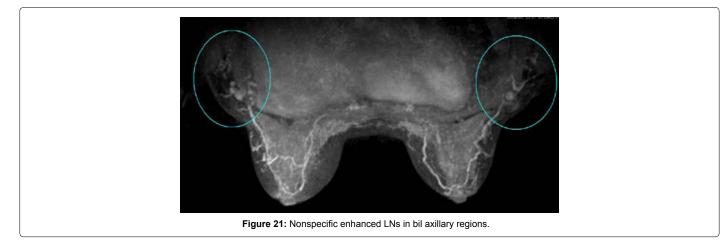


Figure 20: Area of mastitis with impending abscess formations at superior lateral of right breast.



then lead to scarring and calcification. Under mammography, the clusters of microcalcifications in the breast may be observed but cannot be distinguished from malignant calcifications. A common biopsy or a biopsy using mammographically guided needle localization is needed [39,40].

On our dynamic breast MRI, the survival of fat grafted tissues can be qualitatively observed and quantitatively measured. Also, the most frequent finding of complication was focal or diffusely asymmetrical signal intensity changes with fluid or semi-fluid content, without significant mass effect. Nodular lesions, mass-like enhancement, ringlike enhancement, and nodular enhancement were observed. Although, most of the dynamic curves of post-gadolinium enhancement did not have characteristic findings, but based on the morphology and subtracted image, malignancy could be ruled out. From our result, we found four cases that had received operation, drainage or aspiration, but only five cases had higher levels of one or more than one of the following items: CRP, IgE or complement (C4) /ANA. We found the laboratory data are not able to predict all the corresponding post fat grafting infection and rejection with granuloma formations. It supports our contention that breast MRI should acting as first line screening method for early diagnosis and for treatment.

Conclusion

This is a study of using breast MRI to evaluate the conditions of post autologous fat grafting complications. Chronic inflammation with granuloma formations and fat necrosis has the highest incidence rate; we also found acute infection with abscess is the most urgent condition. Overall, 19% of the cases suffering complications had need of an additional operative treatment. Serum test for laboratory data should not be independently used as monitoring parameter for early detection of fat necrosis, granulomatous mastitis or abscess. However, breast MRI is a reliable imaging tool for precise diagnosis on the fat survival, fat necrosis, and other complications.

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Page 12 of 12

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