Magnetic Resonance Imaging and Ultrasound Evaluation after Breast Autologous Fat Grafting Combined with Platelet-Rich Plasma

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Background: Breast lipofilling is a fairly simple and safe procedure if it is performed by experienced surgeons.

Methods: The authors evaluated the radiologic findings from 24 breasts (15 women) subjected to a lipofilling procedure (two sessions) for corrective surgery or cosmetic reasons. Mammography, ultrasound, and magnetic resonance imaging were performed before the first lipofilling session (T0) and 12 months after the last session (T12); ultrasound and magnetic resonance imaging were used 3 months after the first session (Ti) and 3 and 6 months after the last session (T3 and T6). Volumetric evaluations were also made through three-dimensional magnetic resonance imaging reconstruction.

Results: Ultrasound showed oily cysts in 66.67 percent of the breasts at Ti, 70.83 percent at T3, 62.5 percent at T6, and 45.83 percent at T12, whereas magnetic resonance imaging detected oily cysts in 8.33 percent at Ti and T3 and T6 months and 4.17 percent at T12. At Ti, T3, and T6, the cytosteatone-crotic areas identified on both ultrasound and magnetic resonance imaging were unchanged (8.33 percent), whereas at T12 those cytosteatonecrotic areas were increased on ultrasound (12.5 percent) and even more on the magnetic resonance imaging scans (16.67 percent). The average resorption percentage of injected volume was 15.36 percent at T6 months and 28.23 percent at T12 months.

Conclusions: Postlipofilling breast changes can be distinguished from malignant alterations by experienced radiologists and need not interfere with early cancer diagnosis if patients are checked regularly. Moreover, magnetic resonance imaging is very useful for breast volume assessments and for detecting possible changes during longitudinal study. (*Plast. Reconstr. Surg.* 132: 498e, 2013.)

CLINICAL QUESTION/LEVEL OF EVIDENCE: Therapeutic, IV.



any procedures of "oncoplastic" surgery have been developed recently. They combine the radicality of oncologic surgery (wide resection) with plastic surgery techniques, and allow for adequate aesthetic results even where there is an unfavorable ratio between tumor size and gland dimensions. ¹⁻³ Moreover, these procedures are less expensive.

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In particular, lipofilling has awakened a great interest for its considerable advantages: outpatient procedure, easy availability of donor tissue, absence of scar and complications related to implants, and the possibility to change the results obtained by repeating the procedure.⁴ Moreover, apart from lipofilling, no other technique has given such satisfactory results in correcting breast-conserving surgery results.⁵ However, this technique has been criticized, either because a certain degree of fat resorption may occur or because, according to some authors, it can result in breast changes that can make the early diagnosis of cancer difficult.⁴

Disclosure: The authors have no financial interest to declare in relation to the content of this article.

| Patient | Age (years) | Breast | Pre-lipofilling anamnesis | RT | BMI (Kg/m²) | | |
|---------|----------------|--------|--|-----------------------------------|-----------------------|----|-------|
| 01 | 38 | right | BCS with residual scar | yes | 23,15 | | |
| 02 | 74 | right | BCS, prosthesis and retracted scar | yes | 24.77 | | |
| | /4 | left | Mastopexy | no | 24,77 | | |
| 03 | 63 | right | ht Mastectomy and reconstruction with prosthesis | | 24,65 | | |
| | · | left | Breast augmentation with prosthesis | no | | | |
| 04 | 52 | right | Breast augmentation with prosthesis | no | 22,03 | | |
| 04 | | left | Breast augmentation with prosthesis | t augmentation with prosthesis no | | | |
| 05 | 63 | right | Mastectomy and reconstruction with prosthesis | yes | 22.80 | | |
| | | left | Mastactomy and reconstruction with | | 22,89 | | |
| 06 | 68 | right | Breast augmentation with prosthesis | no | 21.26 | | |
| | | left | Breast augmentation with prosthesis | no | 21,36 | | |
| 07 | 55 | right | BCS with residual scar | yes | 24,83 | | |
| 07 | | left | BCS with residual scar | no | | | |
| 08 | 43 | right | BCS with residual scar | no | 22.50 | | |
| 08 | | left | BCS with residual scar | yes | 22,59 | | |
| 09 | 21 | right | Hypomastia, no surgery | no | 21,97 | | |
| 10 | 51 | right | BCS with residual scar | no | 22,06 | | |
| 11 | 28 | right | Removal of prosthesis for rejection | no 21.3 | 21.2 | | |
| | | left | Removal of prosthesis for rejection | no | 21,3 | | |
| 12 | 22 | 22 | 22 | right | Reduction mammaplasty | no | 22.22 |
| | | left | Prosthesis for hypomastia | no | 23,22 | | |
| 13 | 42 | right | BCS and burn scar due to home accident | no | 20,7 | | |
| 14 | 38 | left | Mastectomy and reconstruction with prosthesis | yes | 22,48 | | |
| 15 | 36 | left | BCS, prosthesis and retracted scar | yes | 18,87 | | |
| | mean: 46,27 | | | | _ | | |

Fig. 1. Patient data. Patient, age, breast selected for lipofilling, prelipofilling anamnesis and treatment, radiotherapy, and body mass index for the 15 patients subjected to autologous fat grafting to correct the results of breast-conserving surgery or mastectomy in breast with previous cancer (*white rows*) or simply for cosmetic reasons (*gray rows*). *RT*, radiotherapy; *BMI*, body mass index; *BCS*, breast-conserving surgery.

In this preliminary study, we evaluated radiologic findings in 24 breasts subjected to lipofilling by mammography, ultrasound, and magnetic resonance imaging. A volumetric evaluation of the breast was also made through three-dimensional magnetic resonance imaging reconstruction.

PATIENTS AND METHODS

Population Study

From September of 2009 to May of 2012, we evaluated the radiologic aspects of 24 breasts of 15 female patients (Fig. 1) who underwent lipofilling

to correct surgery results or solely for cosmetic reasons. In all patients, the procedure was carried out in two sessions.

All patients underwent the following examinations: mammography, ultrasound, and magnetic resonance imaging before the first lipofilling session and 12 months after the last lipofilling session; and ultrasound and magnetic resonance imaging 3 months after the first lipofilling session (before beginning the second) and then 3 and 6 months after the last lipofilling session.

The study received the ethics committee's approval, and after a clear explanation of the

benefits and potential risks, all patients gave their informed consent. They also consented to preoperative and postoperative examinations. They were advised to undergo regular checkups during their lifetime with radiologic imaging.

Autologous Fat Grafting

Before and after each procedure, we performed a careful anamnesis and a clinical examination and took photographs to document improvement or the disappearance of defects (Fig. 2, left). Before the first lipofilling session, mammography, ultrasound, and magnetic resonance imaging were performed to rule out signs of tumor recurrence and to have an initial point of comparison to identify new lesions. Only patients who showed no signs of malignancy were included in this study. All procedures were performed under general anesthesia at least 3 months after surgery and at least 6 months after the end of radiotherapy/chemotherapy. Only one 21-yearold patient underwent lipofilling directly, without prior surgery.

The donor site was chosen based on the patient's natural fat deposition.⁶⁻⁸ Before adipose tissue was harvested, 200 ml of Klein solution was injected into the donor site using a specific

cannula (Coleman Kit, Tucson, Ariz.). 9-11 The adipose tissue was then purified by centrifugation (3000 rpm for 4 minutes) and combined with platelet-rich plasma. It was then reinjected aseptically with a specific microcannula (Coleman Kit), using the drop-to-drop technique in small pulses (0.2 to 1 ml), in a radial retrograde manner, on different planes into multiple areas of the breast. According to the patient's needs, in each of the two session, 50 to 150 ml (average, 93.54 ml) was injected, for a total of 187 ml (range, 110 to 250 ml) per patient.

Mammography

Digital mammographic examinations were performed with the GE Senographe DS system (General Electric, Milwaukee, Wis.). Bilateral mammograms with craniocaudal, mediolateral oblique, and mediolateral views were acquired. If necessary, the examinations were completed with additional projections, details, and mammographic magnification.

Ultrasound

Ultrasound studies were performed using a 7- to 15-MHz transducer (ATL HDI 5000; Philips Healthcare, Best, The Netherlands).

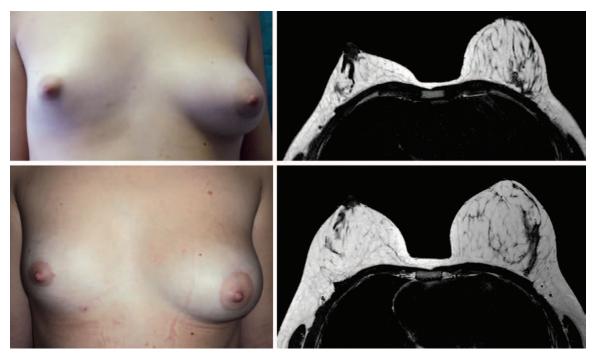


Fig. 2. Photographs (*left*) and axial T2-weighted magnetic resonance imaging scans (*right*) obtained before the first lipofilling session (*above*) and 6 months after the last lipofilling session (*below*), of a 21-year-old patient, subject to no prior surgery, on whom the lipofilling procedure was performed to correct right hypomastia. Lipofilling was carried out on the right breast only, where there was an obvious increase in volume at 6 months after the last lipofilling session compared with before the first lipofilling session.

Magnetic Resonance Imaging

Dynamic magnetic resonance imaging was performed with a 1.5-T (Intera; Philips Healthcare) magnetic resonance imaging apparatus with SENSE technology between days 7 and 14 of the menstrual cycle in fertile women. The patient was placed in prone position with both breasts inside the dedicated bilateral breast surface coil, to avoid any compression. The images were acquired on axial and sagittal planes (Figs. 2 and 3). The magnetic resonance imaging protocol consisted of the following: T1-turbo spin echo (repetition time, 6.8 msec; echo time, 3.3 msec; thickness, 3 mm; gap, 0), T2-turbo spin echo (repetition time, 3800 msec; echo time, 140 msec; thickness, 3 mm; gap, 0), three-dimensional T2-turbo spin echo (repetition time, 2000 msec; echo time, 200 msec; voxel size: anteroposterior, 1 mm; right-to-left, 1 mm; foot-to-head, 1.5 mm), T2-short tau inversion recovery (repetition time, 4000 msec; echo time, 42 msec; inversion time, 155 msec; thickness, 3.0 mm; gap, 0), and T1 dynamic sequence (two-dimensional) (fast field echo) (repetition time, 290 msec; echo time, 4.6 msec; flip angle, 90 degrees; thickness, 3 mm; 8 dynamics; with 50-second time resolution for each), after gadolinium bolus injection [gadopentetic acid and dimeglumine salt (Magnevist; Schering, Berlin, Germany), 0.1 mM/kg at 2 ml/second] followed by saline solution (20 ml). The presence of any lesions and/or enhancement characteristics was studied. Based on magnetic resonance imaging

scans acquired, volumetric assessments of the breasts were also calculated, taking as edges the anterior axillary line, anterior margin of the pectoral muscle, mediosternal line, skin, and nipple. They were assessed using a three-dimensional reconstruction on a separate workstation (ADW 4.0; GE Medical Systems, Milwaukee, Wis.). Finally, at 6 and 12 months after the last lipofilling session, the resorption percentage for each breast was evaluated. Then, we calculated their overall average, and the average for the breasts with oily cyst resorption and for those remaining without resorption. All examinations were performed and analyzed in a blinded fashion by two radiologists experienced in interpreting breast imaging.

Statistical Analysis

The McNemar test was performed to compare ultrasound and magnetic resonance imaging in finding oily cysts and cytosteatonecrosis at 3 months after the first lipofilling session and at 3, 6, and 12 months after the last lipofilling session. A value of p < 0.05 was considered statistically significant. Because of the small sample size of this preliminary study, a statistical analysis on volumetric data was not feasible.

RESULTS

Radiologic Findings

No pathologic alterations were found in any of the examined breasts on mammography,

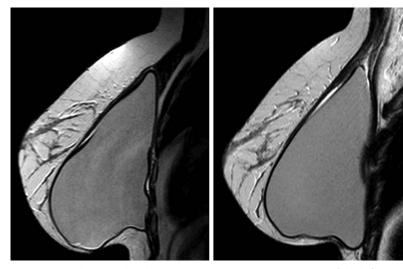


Fig. 3. Sagittal T2-weighted magnetic resonance imaging scans of the left breast obtained before the first lipofilling session (*left*) and 3 months after the first lipofilling session (*right*), of a 22-year-old patient on whom the lipofilling procedure was performed, to correct left hypomastia, after an augmentation mammaplasty with prosthesis.

Table 1. Findings from Radiologic Examinations

| Findings | Ti (%) | T3 (%) | T6 (%) | T12 (%) |
|-----------------------------|---------------|---------------|--------------|---------------|
| Mx | | | | |
| Microcalcification | _ | _ | _ | 5/24 (20.83) |
| Macrocalcification | _ | _ | _ | 3/24 (12.5) |
| Liponecrosis | _ | _ | _ | 3/24 (12.5) |
| US | | | | |
| Oily cysts (5–10 mm) | 16/24 (66.67) | 17/24 (70.83) | 15/24 (62.5) | 11/24 (45.83) |
| Cytosteatonecrosis (>20 mm) | 2/24 (8.33) | 2/24 (8.33) | 2/24 (8.33) | 3/24 (12.5) |
| MRÍ | | | | |
| Oily cysts | 2/24 (8.33) | 2/24 (8.33) | 2/24 (8.33) | 1/24(4.17) |
| Cytosteatonecrosis | 2/24 (8.33) | 2/24 (8.33) | 2/24 (8.33) | 4/24 (16.67) |

Ti, 3 months after the first lipofilling session; T3, 3 months after the last lipofilling session; T6, 6 months after the last lipofilling session; T12, 12 months after the last lipofilling session; Mx, mammography; US, ultrasound; MRI, magnetic resonance imaging.

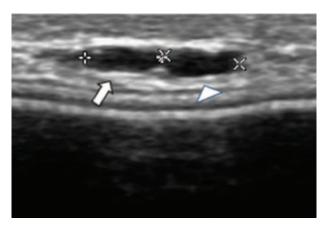


Fig. 4. Ultrasound image of the right breast of a 22-year-old patient that shows two oily cysts, with oval hypoechoic/anechoic areas with a well-defined regular wall and sizes of 4.6 (*arrow*) and 4.5 mm (*arrowhead*), respectively.

ultrasound, or magnetic resonance imaging before the first lipofilling session. In 23 breasts, the previous surgery outcomes were visible. With regard to postlipofilling examinations, we report below only those findings that were not present before the lipofilling (Table 1).

ultrasound, On oily cysts (round hypoechoic/anechoic areas with a well-defined regular wall and sizes up to 10 mm) (Figs. 4 and 5) were found in 66.67 percent of the breasts at 3 months after the first lipofilling session, in 70.83 percent at 3 months after the last lipofilling session, in 62.5 percent at 6 months after the last lipofilling session (showing oily cyst resorption in two breasts), and in 45.83 percent at 12 months after the last lipofilling session (showing oily cyst resorption in four other breasts); whereas on magnetic resonance imaging, they were detected in 8.33 percent at 3 months after the first lipofilling session and 3 and 6 months after the last lipofilling session, and in 4.17 percent at 12 months after the last lipofilling session (showing oily cyst resorption

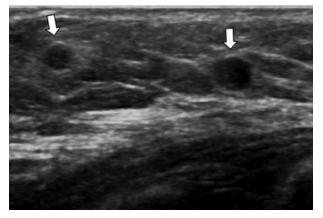


Fig. 5. Ultrasound image of the right breast of a 42-year-old patient that shows oily cysts (*arrows*), with round hypoechoic/anechoic areas with a well-defined regular wall and sizes ranging between 5 and 10 mm.

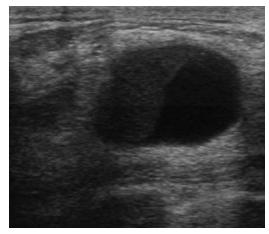
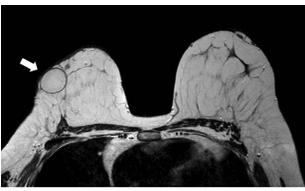


Fig. 6. Ultrasound image of the right breast of a 38-year-old patient that shows a cytosteatonecrotic area (hypoechoic/anechoic area) of approximately 4 cm.

in only one breast). At 3 months after the first lipofilling session and 3 and 6 months after the last lipofilling session, the cytosteatonecrotic areas (anechoic areas, >20 mm) (Figs. 6 and 7)



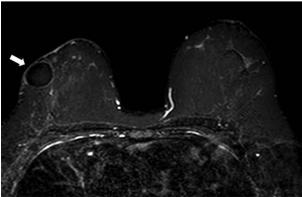


Fig. 7. Axial T2-weighted (*above*) and short tau inversion recovery (*below*) magnetic resonance imaging scans of the breasts obtained at 6 months after the last lipofilling session, of a 55-year-old patient on whom the lipofilling procedure was performed to correct bilateral scarring. On the right breast, a cytosteatonecrotic area clearly appears: a round area of approximately 28 mm, hyperintense on T2-weighted images (*arrow*), and the signal of which is suppressed on short tau inversion recovery sequences (*arrow*).

identified on both ultrasound and magnetic resonance imaging were unchanged (8.33 percent), whereas at 12 months after the last lipofilling session the breasts with cytosteatone-crotic areas were increased on ultrasound (12.5 percent) and even more on magnetic resonance imaging (16.67 percent).

At 12 months after the last lipofilling session, the mammograms showed microcalcifications in 20.83 percent: in three cases, they had typical benign features (scattered and under various guises), and in two cases, they are clustered. Six months later, the latter were subjected to follow-up mammography and reclassified as benign. Macrocalcifications were found in 12.5 percent and fat necrosis (areas of radiolucency with a thin, dense, regular rim) was found in 12.5 percent. Subsequent to the lipofilling procedure, no solid mass or opacity was found in any patient.

Volumetric Assessment at 6 and 12 Months after the Last Lipofilling Session

At 6 months after the last lipofilling session, the average resorption of the injected volume was 15.36 ± 1.76 percent (range, 9.95 to 17.55 percent) for all the breasts and 14.40 ± 0.68 percent (range, 13.92 to 14.88 percent) for only the two breasts with oily cyst resorption and 15.94 ± 1.31 percent (range, 13.88 to 17.55 percent) for only the breasts without oily cyst resorption (Fig. 8). At 12 months after the last lipofilling session, the average resorption was 28.23 ± 1.55 percent (range, 25.47 to 31.99 percent) for all the breasts, 27.84 ± 1.15 percent (range, 26.72 to 29.88 percent) for the six breasts with oily cyst resorption, and 28.23 ± 1.96 percent (range, 25.47 to 31.99 percent) for the breasts without oily cyst resorption.

Statistical Analysis

Using the McNemar test, we found a statistically significant (p < 0.05) difference between the oily cysts detected with ultrasound and those identified with magnetic resonance imaging, at any time (Fig. 9). However, we have not found a statistically significant difference for the cytosteatonecrotic areas (Fig. 10).

DISCUSSION

Autologous fat transplantation was initially performed by Neuber to fill a depressed scar.¹² Since then, some plastic surgeons have applied this procedure in almost all body regions.⁸ Breast lipofilling should be performed only by well-trained and skilled surgeons to avoid major complications, which are mainly attributable to technical errors.^{13–15}

One of the main reasons why this technique was questioned is that there may be lipofilling resorption. Therefore, the results are unpredictable. 7,16–18 In the literature, the resorption rate reported over the first year is highly variable (20 to 90 percent), most evidently between the fourth and sixth months.^{7,18–24} However, so far, in many studies, the evidence of breast lipofilling survival was based on patient satisfaction and plastic surgeons' evaluations. 10,17,25-29 In a study where mammary volumes were calculated by computed tomography using a three-dimensional program, a resorption rate of 47.5 percent at 9 months was reported.²⁶ However, computed tomography is not indicated for longitudinal studies because it can cause tumors induced by radiation. Instead, magnetic resonance imaging allows for a good volume estimate^{20,30} and does not pose this risk.

| PATIENTS | BREAST | INITIAL BREAST VOLUME (ml) | INJECTED VOLUME (ml) | BREAST VOLUME AT T ₆ . | RESORPTION % AT T ₆ . | BREAST VOLUME AT T ₁₂ . | RESORPTION % AT T ₁₂ . |
|----------|--------|----------------------------------|-------------------------|---|----------------------------------|--|-----------------------------------|
| 01 | right | 364,59 | 120 | 463,60 | 17,49 | 448,15 | 30,37 |
| 02 | right | 713,18 | 180 | 861,87 | 17,39 | 835,59 | 31,99 |
| | left | 810,50 | 110 | 901,20 | 17,55 | 887,56 | 29,94 |
| 03 | right | 812,68 | 220 | 996,83 | 16,30 | 967,25 | 29,74 |
| | left | 1078,29 | 180 | 1230,02 | 15,71 | 1209,00 | 27,38 |
| 04 | right | 1063,46 | 200 | 1231,98 | 15,74 | 1208,00 | 27,73 |
| | left | 965,99 | 250 | 1173,81 | 16,87 | 1148,22 | 27.11 |
| 05 | right | 612,30 | 200 | 780,56 | 15,87 | 758,12 | 27.09 |
| | left | 690,88 | 200 | 858,72 | 16,08 | 831,12 | 29,88 |
| 06 | right | 1003,53 | 200 | 1174,23 | 14,65 | 1145,36 | 29,08 |
| | left | 1072,28 | 200 | 1238,74 | 16,77 | 1215,35 | 28,46 |
| 07 | right | 754,64 | 200 | 926,44 | 14,10 | 900,22 | 27,21 |
| | left | 811,59 | 200 | 978,53 | 16,53 | 957,74 | 26,92 |
| 08 | right | 560,00 | 200 | 726,64 | 16,68 | 702,55 | 28,72 |
| | left | 541,92 | 200 | 713,42 | 14,25 | 690.98 | 25,47 |
| 09 | right | 367,18 | 180 | 521,98 | 14,00 | 498,11 | 27,26 |
| 10 | right | 247,44 | 200 | 417,68 | 14,88 | 394,00 | 26,72 |
| 11 | right | 198,89 | 200 | 365,57 | 16,66 | 342,87 | 28,01 |
| | left | 226,53 | 200 | 395,03 | 15,75 | 371,85 | 27,34 |
| 12 | right | 1056,57 | 150 | 1187,58 | 12,66 | 1161,64 | 29,95 |
| | left | 1020,62 | 150 | 1149,74 | 13,92 | 1128,06 | 28,37 |
| 13 | left | 311,64 | 150 | 446,72 | 9,95 | 418,11 | 29.02 |
| 14 | left | 799,82 | 200 | 972,06 | 13,88 | 948,12 | 25,85 |
| 15 | right | 593,42 | 200 | 763,56 | 14,93 | 737,58 | 27,92 |

Fig. 8. Transplanted fat resorption percentage 6 and 12 months after the last lipofilling session. *Light gray*, patients with oily cysts; *dark gray*, patients with oily cyst resorption; *Injected Volume*, total fat volume injected in the two lipofilling sessions; $T_{6'}$ 6 months after the last lipofilling session; $T_{12'}$ 12 months after the last lipofilling session.

Serial magnetic resonance imaging offers a quantitative measure of fat resorption and survival. 31,32 We found a total average resorption percentage of 15.36 percent at 6 months after the last lipofilling session and of 28.23 percent at 12 months after the last lipofilling session. Evaluation at 3 months after the last lipofilling session may be too early and incomplete.

To prevent, or rather minimize, resorption, it is crucial to perform each step of the procedure carefully, paying close attention to the technical details. We injected 200 ml of Klein solution into the donor site before liposuction. The lipoaspirate must then be purified in various ways: washed with different solutions, ^{19,33–35} filtered, centrifuged, and left to decant. ^{15,36} Its injection causes an inflammatory response that can be reduced by injecting only adipocytes. ^{37,38} An inverse relationship has been observed between the blood amount in the lipoaspirate and the viable number of adipocytes. ^{39,40} Furthermore, the fat must be in contact

with air as little as possible, 19 as this may cause lysis of the adipocytes. $^{36-41}$

In our patients, the lipoaspirate was purified by centrifugation and combined with platelet-rich plasma. Platelet-rich plasma has no impact on the diagnostic images but improves lipofilling results and reduces the resorption rate, increasing fatgraft survival. 42 Then, the lipoaspirate was injected using the drop-to-drop technique and in multiple sessions to maximize the contact surface between the lipoaspirate and the host's capillaries.¹⁸ Diffusion of nutrients from neighboring capillaries is essential for adipocyte survival and favors their integration with the surrounding tissue. 15,24,25 The need for multiple sessions of extended treatment is the main factor that discourages patients. 15 Finally, the problem of fat resorption may be resolved by injecting a total volume of fat greater than the desired volume.^{7,19}

Other possible lipofilling complications are swelling, bruising, infection, granuloma or abscess,

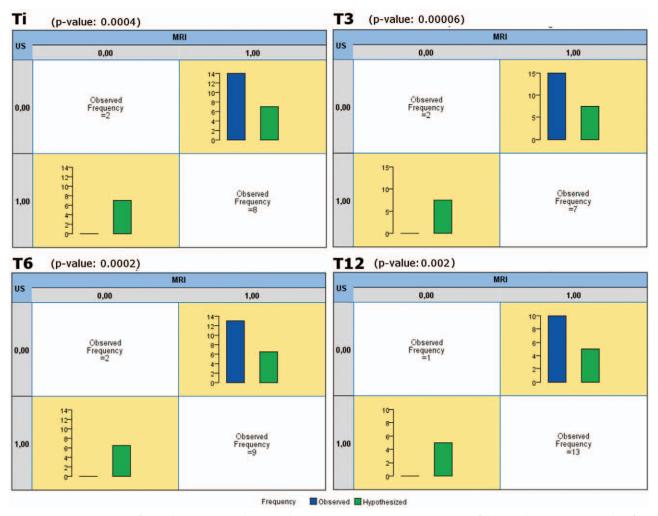


Fig. 9. McNemar tests performed to compare ultrasound and magnetic resonance imaging in finding oily cysts at 3 months after the first lipofilling session (Ti) and at 3 (T3), 6 (T6), and (T12) 12 months after the last lipofilling session. A value of p < 0.05 was considered statistically significant. We found a statistically significant (p < 0.05) difference between the oily cysts detected with ultrasound and those identified with magnetic resonance imaging, at any time.

simple or complicated cysts, fat necrosis, calcification, masses, and rarely nipple retraction caused by fat necrosis. 4,8,43 Microcalcification and liponecrosis are the main reason why this technique was abandoned at the end of the 1980s. It was thought that lipofilling could make the early diagnosis of eventual underlying cancers difficult.^{5,6} This is open for discussion. First, liponecrosis, oily cysts, calcification, and focal opacity can be caused by any type of breast surgery (biopsy, reduction mammoplasty, 44,45 breast reconstruction, 46 liposuction⁴⁷) or can occur spontaneously.^{46,48,49} In a retrospective study, calcifications were found in 50 percent of cases after reduction mammaplasty.⁵⁰ However, they were not held responsible for interfering with early cancer diagnosis and the procedure was not abandoned.^{5,6} We found microcalcifications in 20.83 percent, the same value

observed in a study where the postlipomodeling incidence of microcalcification was equal to that found following surgery (20 percent), for a total of 40 percent.⁵ These data suggest that microcalcifications can be a normal consequence of lipomodeling. Usually, they are easily identified (like the ones formed after breast surgery) because of their typical benign radiologic features (small, round, and regular; isolated or associated with small radiolucent images of lipidic nature, scattered or more rarely clustered; usually classified as Breast Imaging-Reporting and Data System 2), and they are very different from the ones suspected of malignancy or recurrence (because of localization or radiologic aspects). 5,44,46,51-53 Calcifications suspected of being malignant were found in none of our patients. In three breasts, microcalcifications appeared to have typical benign

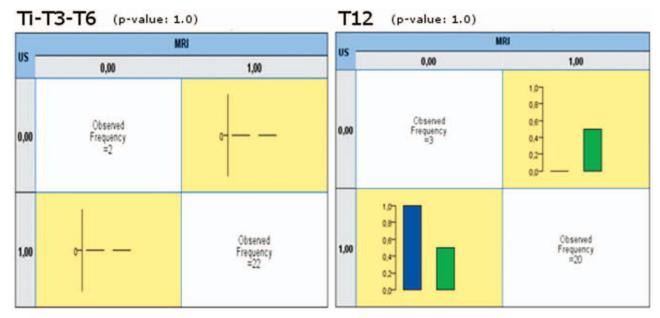


Fig. 10. McNemar tests performed to compare ultrasound and magnetic resonance imaging in finding cytosteatonecrosis at 3 months after the first lipofilling session (Ti) and at 3 (T3), 6 (T6), and (T12) 12 months after the last lipofilling session. A value of p < 0.05 was considered statistically significant. We have not found a statistically significant difference between the cytosteatonecrotic areas detected with ultrasound and those identified with magnetic resonance imaging.

features. In two other cases, microcalcifications were clustered, although the subsequent control reclassified them as benign.

Oily cysts and cytosteatonecrotic areas are manifestations of liponecrosis. Therefore, it is important to understand their etiopathogenesis, both to prevent them and to distinguish their



Fig. 11. Mammogram of a 51-year-old patient that clearly shows the typical soap bubble feature: a radiolucent lesion confined by a thin calcified rim (*arrow*).

manifestations from malignancies. The etiopathogenesis of liponecrosis seems to be complex and multifactorial. Besides surgery, the most common causes of fat necrosis are ischemia, radiotherapy, trauma, and sometimes anticoagulant therapy. 49,53-60 Postlipofilling liponecrosis has been correlated to mechanical trauma caused by the blunt cannula used for fat injection and to a higher tendency to fat graft necrosis.⁵³ According to others, liponecrosis is caused by an inadequate blood supply of the transplanted fat. Adipocyte suffering results in lysis, releasing fatty acids and glycerol, which may affect the osmotic balance and promote edema, thus worsening the capillary perfusion. Then, there is an inflammatory response of the host tissue to the degenerated fat, subsequently replaced by fibrosis.^{25,61} When a peripheral fibrosis ("capsule") develops, cysts are often visible only with ultrasound (Figs. 4 and 5).25,53 After a few months, the subtle fibrotic capsule can thicken and calcify; therefore, cysts become visible also on mammography, sometimes with the typical "soapbubble" feature (a radiolucent lesion confined by a thin calcified rim) (Fig. 11). Furthermore, in the liponecrotic areas, microcalcifications or macrocalcifications (visible with mammography) can form (Fig. 12).⁶² However, the fibrosis can mostly extend, forming a spiculated mass, that can be suspected of being malignant.⁵³ In our patients, no solid masses were found.



Fig. 12. Lateral (*above*) and mediolateral oblique (*below*) bilateral mammograms of a 43-year-old patient that show calcifications of the liponecrotic areas.

If patients are checked regularly, experienced radiologists are able to identify postlipofilling breast changes and distinguish them from malignant alterations so that lipofilling does not make early diagnosis of cancer difficult. Although there is still no postlipomodeling screening protocol,⁵ it is important to perform closer checks during the first year (e.g., ultrasound and magnetic resonance imaging at 3, 6, and 12 months after the last lipofilling session, and mammography at 12 months after the last lipofilling session) and follow-up annually, even in the absence of alterations, especially in patients who have already had cancer. Mammography, ultrasound, and magnetic resonance imaging before the first lipofilling session are necessary

to rule out any signs of tumor recurrence and to have an initial point of comparison, whereas at 3 months after the first lipofilling session, the examinations do not seem to add important data.

In patients who have had cancers or are aged 35 years and older, it is important to perform prelipofilling mammography and follow-up annually or perform closer checks (the frequency of which depends on the type of alteration found). Mammography, though still the criterion standard,^{53,63} may result in false-positives (18 percent). 64,65 Mammography detects only two-thirds of recurrences, and in approximately 45 percent of cases, the recurrences (in situ or invasive) occur as microcalcifications.⁶³ Mammography is the only examination that can clearly identify microcalcifications and distinguish them from those that are malignant.⁵ In addition, mammography allows us to identify the typical soap bubble, macrocalcifications, and breast opacity. It is therefore a key examination in postlipofilling monitoring; however, alone, it is not enough. 5,53,62 In patients younger than 35 years who have undergone lipofilling solely for aesthetic reasons, mammography is not indicated.

Ultrasound is more reliable than magnetic resonance imaging for detecting small cysts (thus, it is more sensitive), but it cannot distinguish the oily cyst from the normal cyst (less specific). The small oily cysts are best seen by ultrasound, because they behave like small fluid formations. In our patients, small oily cysts appeared on a much higher percentage of breast ultrasound images as opposed to their magnetic resonance imaging scans (as confirmed by the McNemar test), highlighting also their reabsorption (Table 1). However, the oily cysts cannot be differentiated from normal cysts on ultrasound. In one of our patients, normal cysts mistaken at first for oily cysts on ultrasound were correctly identified through the subsequent magnetic resonance imaging scan.

Ultrasound is also extremely useful in the evaluation of "dense breasts,"⁶⁶ surgical scars and complex cysts, and in the differentiation between cysts and solid masses. Finally, when in doubt, it can be used to perform ultrasound-guided microbiopsy.⁵

The role of magnetic resonance imaging, although emphasized in recent studies, is not yet clear. In magnetic resonance imaging, oily cyst signal intensity is suppressed in sequences with fat suppression because they are composed of fat even if necrotic. However, small oily cysts cannot be seen properly because of their small size, the result of which is that they are easily confused with native fat (also suppressed). In those two breasts where the oily cysts were visible also on magnetic

resonance imaging, the oily cysts were larger compared with those of other patients. Even though magnetic resonance imaging does not always detect small oily cysts, it can positively differentiate the normal cysts because they are hypointense on T1-weighted sequences, hyperintense on T2-weighted sequences, and not suppressed on the sequences with fat suppression. Moreover, because of the sequences where fat suppression occurs, magnetic resonance imaging is able to highlight large cytosteatonecrotic areas. Magnetic resonance imaging seems to be at least as effective as ultrasound and mammography in demonstrating large cytosteatonecrotic areas, as confirmed by the McNemar test. However, these data need to be confirmed using a wider test population.

In addition, magnetic resonance imaging is extremely useful in cases of suspected recurrence, namely, clinical signs of recurrence together with negative or equivocal standard radiologic examinations, and scar alteration. In fact, it has excellent sensitivity and specificity (>90 percent) in the diagnosis of infiltrating recurrence, but because of large variation in angiogenesis, it remains limited in pure intraductal forms (sensitivity between 60 and 85 percent).67 Dynamic studies with contrast agents, fat saturation, and image subtraction improve the specificity of breast magnetic resonance imaging.⁵ It would be optimal to perform magnetic resonance imaging at least 1 year after radiotherapy because, as a result of inflammation and posttreatment edema, there may be benign enhancements.66

Furthermore, magnetic resonance imaging can be used for breast volume assessment and for possible changes in longitudinal studies.^{20,30} Thus, it can be very useful for better understanding the behavior of transplanted fat and the mechanisms that underlie its reabsorption.

In some patients, probably because of resorption, we found a reduction of oily cysts over time (in number and size). In these patients, the fat resorption percentage was similar to that of others. However, in these patients, a better aesthetic result was obtained. These data need confirmation and further investigation on a wider test population.

CONCLUSIONS

In addition to ultrasound and mammography, magnetic resonance imaging is very useful for evaluation of medium- and long-term complications. It is also very useful for a quantitative breast-volume assessment and provides early information on fat resorption.

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