



Combined microvascular breast and lymphatic reconstruction with deep inferior epigastric perforator flap and gastroepiploic vascularized lymph node transfer for postmastectomy lymphedema patients

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Background: The combination of microvascular breast reconstruction (MBR) and vascularized lymph node transfer (VLNT) in a single-stage procedure is a surgical option for women who desire breast reconstruction and postmastectomy lymphedema surgery. In this study, we present a series of patients who underwent simultaneous lymphatic and MBR with the gastroepiploic VLNT (GE-VLNT) and the deep inferior epigastric perforator (DIEP) flap respectively.

Methods: Between 2018 and 2019, all consecutive patients diagnosed with lymphedema stage IIB–III International Society of Lymphology who opted to pursue simultaneous MBR with DIEP flap and GE-VLNT were included in this study. Patient demographics, comorbidities, prior radiation therapy, operative characteristics, surgical outcomes and complications were collected and analyzed.

Results: Six patients underwent simultaneous unilateral MBR with DIEP flap and GE-VLNT. The mean age was 48 ± 10.5 years and mean body mass index was 28.2 ± 4.5 kg/m². The flap survival rate was 100%. One patient required re-exploration due to venous congestion of the lymph node flap but was successfully salvaged. There was no donor site morbidity at the donor or recipient site for the DIEP flap were seen. The mean circumference reduction rate was $30.0\% \pm 5.1\%$ ($P < 0.001$). One patient stage III underwent additional liposuction at 12 months postoperative to improve final results.

Conclusions: The combined use of DIEP flap and GE-VLNT flaps in a single-stage procedure is a safe and reliable surgical option for patients with postmastectomy lymphedema who desire and are suitable for autologous microvascular breast and lymphatic reconstruction.

Keywords: Lymphedema; vascularized lymph node transfer (VLNT); breast reconstruction and lymph node transfer; deep inferior epigastric perforator flap and lymph node transfer (DIEP flap and lymph node transfer); gastroepiploic lymph node transfer

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Introduction

Upper extremity lymphedema (UEL) is a complex condition commonly related to breast cancer and its associated oncologic therapies, including cancer resection, radiation and chemotherapy. In the United States, breast cancer is the most common cancer diagnosis in women excluding cutaneous malignancies. Approximately 268,600 women are expected to be newly diagnosed with invasive breast cancer in 2019, and about 41,760 women in the United States are estimated to die from this cancer in the same year (1). Reported incidence of UEL among observational studies varies widely from 7.6% to 49% but is trending down due to less invasive cancer treatment options (2).

The mainstay of treatment for extremity lymphedema is complex decongestive therapy (CDT), which consists of an intensive program that combines bandaging, compression garments, exercise, manual lymphatic drainage, and self-care measurements. If this fails, physiological or debulking surgical procedures are considered. The former attempts to restore or improve the lymphatic flow with procedures such as lymphatico-venous anastomoses (LVA) and vascularized lymph node transfer (VLNT) (3-9). On the other hand, the latter removes the diseased tissue through radical excision or suction-assisted lipectomy (SAL) (3). Both non-surgical and surgical treatment options aim to reduce the circumference of the affected extremity in addition to improve the patient's symptomology, extremity-related morbidities such as skin infections, and quality of life (10).

VLNTs have shown to be successful in treating patients with moderate to advanced stages of lymphedema (6,7,9, 11-16). Two possible theories have been described explaining its mechanism: the "lymphatic wick", which suggests the regeneration of spontaneous connections with efferent lymphatic, and the "lymphatic pump", which proposes an intrinsic lympho-venous shunt within the transferred lymph nodes (17).

Generally, breast reconstruction options encompass the use of tissue expanders/implants or autologous reconstruction based on abdominal, back, gluteal, or thigh flaps. For autologous breast reconstruction, the free deep inferior epigastric perforator (DIEP) flap has increasingly gained popularity due to its versatility, ample volume and comparative advantages over other flaps (18). A large study using the Mastectomy Reconstruction Outcomes Consortium (MROC) showed patient satisfaction with breasts and psychosocial and sexual well-being two years after surgeries were higher with autologous breast

reconstruction (19). However, the odds of complications appear to be higher in this patient group (20).

Autologous breast reconstruction can be simultaneously performed with VLNTs in selected patients, requiring both breast reconstruction and surgical treatment for lymphedema (21-27). The most common vascularized lymph node flap used for this combined MBR and lymphatic reconstruction approach is the groin. In this study, we describe a novel combined surgical approach of autologous breast reconstruction with DIEP and gastroepiploic VLNT (GE-VLNT), and we present a case series of patients who underwent this procedure.

Methods

Between January 2018 and January 2019, all consecutive postmastectomy patients who underwent delayed autologous microvascular breast reconstruction (MBR) and were clinically and radiologically diagnosed with lymphedema, stage IIb–III International Society of Lymphology (ILS) (28) were included in this study. All patients underwent autologous breast reconstruction with the DIEP flap and gastroepiploic vascularized lymph node flap transfer by the lead author (PC) in the same operative time. Prior to surgery, the patients were examined by their oncologists, and all were considered cancer free. Patients older than 75 years, patients with prior surgical treatments for lymphedema or patients requiring bilateral breast reconstruction were excluded. Lymphedema was diagnosed preoperatively by thorough clinical evaluation, limb circumference measurements and lymphoscintigraphy, which served as a reference line to assess objectively response to surgery. Patient demographics, comorbidities, prior radiation therapy, operative characteristics, surgical outcomes and complications were prospectively collected and registered after obtaining approval from the local Ethics Committee.

Clinical photographs both pre- and post-operatively during follow-up were also documented. Circumferences of both the edematous extremity and the unaffected counterpart pre and at least 6 months postoperatively were documented to calculate the reduction rate. The anatomic landmarks used to measure the circumference were the following: 10 cm below the elbow or knee, 10 cm above the wrist or ankle, and at the midhand or midfoot. To calculate the circumference reduction rate, we used the mean of the circumferences measured. We defined the percentage of improvement as "reduction rate" comparing the affected

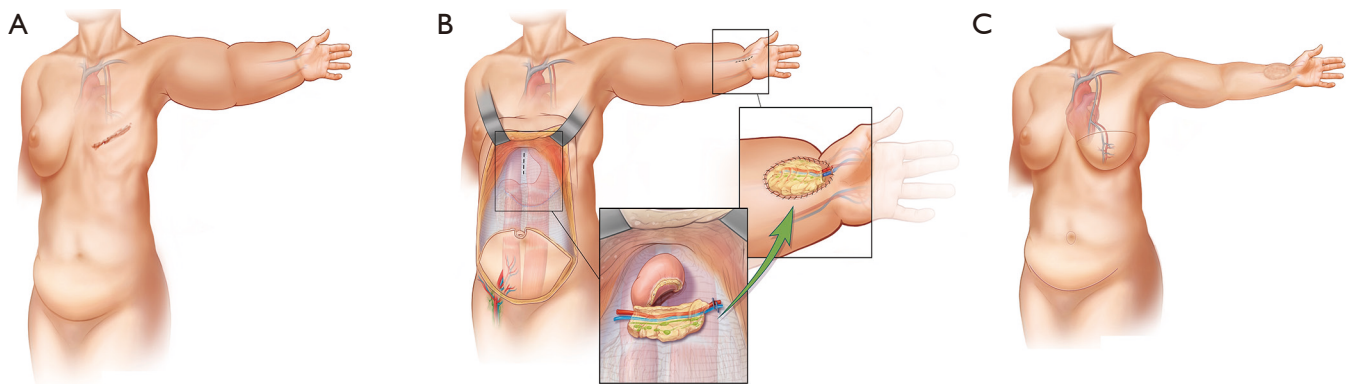


Figure 1 Schematic representation of the combined microvascular breast and lymphatic reconstruction. (A) Post-mastectomy female patient with left breast-cancer related lymphedema (BCRL). (B) The deep inferior epigastric perforator (DIEP) flap harvest. In the upper abdomen, a 6 cm incision is done over the linea alba to access the intra-abdominal space. Part of the stomach and omentum is exposed to harvest the gastroepiploic lymph node flap, which is then transferred to the wrist. (C) Immediate postoperative representation. Notice the reconstructed breast and the limb size reduction.

limb (AL) to the healthy limb (HL) using the following equation:

$$\text{Circumferencereductionrate (\%)} = \frac{[1 - (\text{postoperativeAL} - \text{HL})]}{(\text{pre} - \text{postoperativeAL} - \text{HL})} \times 100$$

Surgical technique

A two-surgical team approach was used. One team dissected and prepared the internal mammary vessels and the palmar branch of the radial artery and cephalic vein for its use as recipient vessels for the DIEP flap and the GE-VLNT, respectively, whereas the other team harvested the flaps. *Figure 1* shows a schematic representation of the combined microvascular breast and lymphatic reconstruction.

Under general anesthesia, the DIEP flap, which we based on the deep inferior epigastric pedicle, was harvested in the standard fashion (29–31). Before dividing the deep inferior epigastric vascular pedicle, attention was then turned to the upper abdomen for the gastroepiploic vascularized lymph node flap harvest (*Figure 2*). The superior abdominal flap was dissected to the level of the xiphoid process. A midline laparotomy incision of about 6 cm was done along the *linea alba*. The greater curvature of the stomach and omentum were identified in the intraabdominal space and then extruded (*Figure 2B*). We cautiously injected 0.5 cc of indocyanine green (ICG) into the omentum and identified the gastroepiploic lymph nodes. The dissection was carried along the greater curvature of the stomach from left to right to isolate the gastroepiploic vessels, and

the flap was harvested, as previously described (13). The proximal portion of the lymph node flap based on the right gastroepiploic vessels were placed in the volar aspect of the wrist and anastomosed to the palmar branch of the radial artery and the cephalic vein. When needed, skin graft was used to cover the flap and close the skin without tension (*Figure 2E*). Extensive scar and fibrotic tissue were removed and released from the axillary vessels before DIEP flap inset. The DIEP flap was anastomosed to the internal mammary vessels. Adequate blood perfusion was evaluated clinically and with the use of Doppler for both flaps.

Postoperative care

All patients were admitted as inpatients for 5 to 7 days postoperatively for flap monitoring, pain control and physical therapy. Drains were removed when the output was less than 30 mL/day. Physiotherapy was started approximately 2 to 3 weeks postoperatively, after complete wound healing and suggested for 3 times a week for the next 6 months postoperatively.

Statistical analysis

Analyses were conducted using JMP[®] version 13 (SAS Institute Inc., Cary, NC, USA, 1989–2019). Measures of central tendency and dispersion were obtained to show data distribution. Wilcoxon signed-rank test was used to evaluate

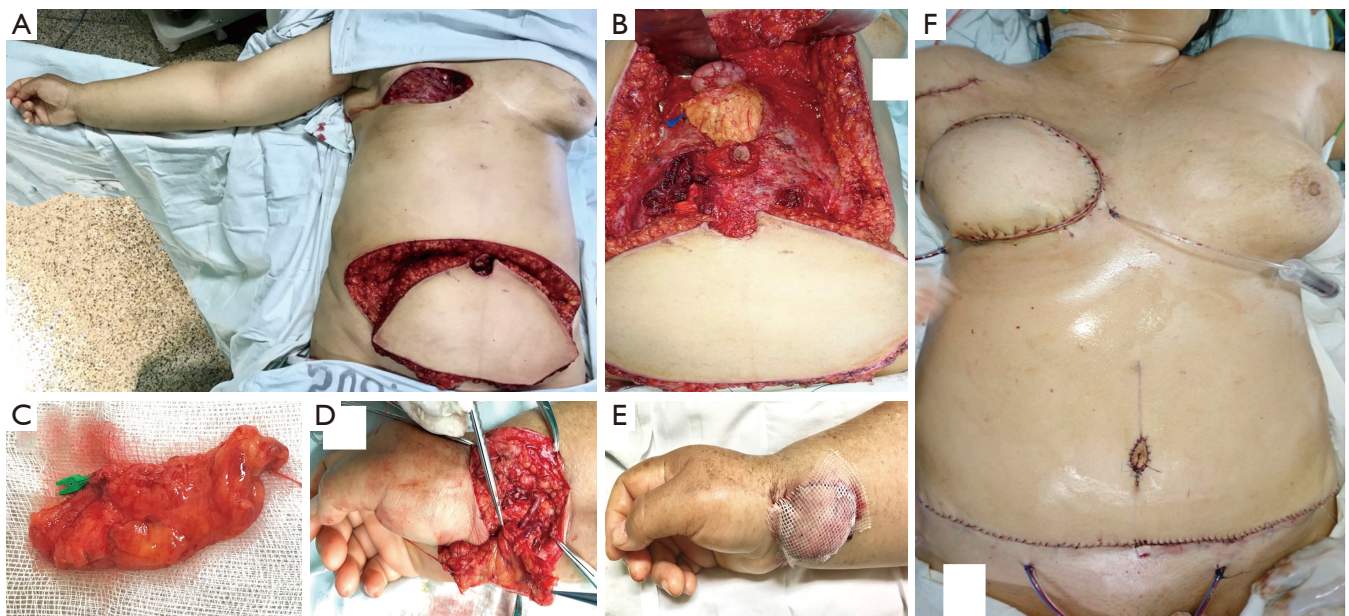


Figure 2 Intraoperative clinical pictures. (A) The deep inferior epigastric perforator (DIEP) flap harvest. (B) Mini-laparotomy performed after the elevation of the upper abdominal flap. Notice part of the stomach and omentum is extruded to harvest the gastroepiploic vascularized lymph node flap. (C) Transferred lymph node flap on the table. (D,E) Inset of the gastroepiploic lymph node flap at the level of the wrist and its closure with a split-thickness skin graft to avoid closure with tension. (F) Immediate postoperative picture after the combined microvascular breast reconstruction with DIEP flap and lymphatic reconstruction with GE-VLNT. GE-VLNT, gastroepiploic vascularized lymph node transfer.

pre and postoperative circumference measurements. Statistical significance level was set at $P < 0.05$.

Results

Six consecutive patients with refractory UEL related to breast cancer treatment underwent simultaneous MBR with DIEP flap and GE-VLNT. The mean age was 48 ± 10.5 years and mean body mass index was 28.2 ± 4.5 kg/m² (Table 1). Five patients (83.3%) had received radiation therapy before surgery. The mean duration of lymphedema symptom was 18.6 ± 5.3 months. All patients underwent CDT for the treatment of extremity lymphedema with a mean duration of 15 ± 1.3 months before surgery.

For all patients, the internal mammary vessels were utilized as recipient vessels for the DIEP flap. In addition, the palmar branch of the radial artery and the cephalic vein at the level of the wrist were used as recipient vessels for the GE-VLNT in all patients. Patients were hospitalized for a mean duration of 15.8 ± 2.5 days. Mean follow-up time was 12.8 ± 2.5 months. Operative time is shown in Table 2.

The mean circumference reduction rate was

$30.0\% \pm 5.1\%$ ($P < 0.001$). All patients required contralateral breast reduction for symmetry. One patient required re-exploration of the lymph node flap due to venous congestion secondary to kinking of pedicle vessels, but was successfully salvaged in the operating room redoing the anastomosis. There were no flap losses in our case series. No major complications were seen at the donor or recipient sites of the DIEP flap. One patient had small skin wound dehiscence in the abdominal donor site. All patients reported subjective symptomatic improvement. One patient with stage III lymphedema underwent additional liposuction of the affected extremity at 12 months postoperative to improve surgical outcomes (Table 3). Figures 3 and 4 show pre and postoperative pictures of two patients who underwent simultaneous autologous MBR and GE-VLNT.

Discussion

Surgical treatment for lymphedema has experienced a rapid growth over the past few years largely attributable to not only the increased awareness of this condition among patients and medical community, but also due to

Table 1 Patient characteristics

Characteristic	Mean \pm SD/count	Range
Age, years	48 \pm 10.5	37.0–61.0
Body mass index, kg/m ²	28.2 \pm 4.5	23.7–31.0
Duration of lymphedema-associated symptoms (months)	18.6 \pm 5.3	8.0–35.0
Duration of preoperative complex decongestive therapy (months)	15 \pm 1.3	12–17
Prior radiation therapy		
Yes	5	
No	1	

SD, standard deviation.

Table 2 Operative data

Variable	Mean \pm SD/count	Range
Hospitalization (days)	15.8 \pm 2.5	12.5–18.0
Follow up time (months)	12.8 \pm 2.5	6.0–18.5
Operative time (min)		
DIEP harvest	105.5 \pm 15	90–115
GE-VLN flap harvest	35 \pm 5.6	30.0–40.5
Total operative time	325 \pm 25.5	300.5–350.5
DIEP Flap weight (gram)	380.4 \pm 62.3	352–440

SD, standard deviation; DIEP, deep inferior epigastric perforator; GE-VLN, gastroepiploic vascularized lymph nodes.

Table 3 Surgical outcomes and complications

Variable	Mean \pm SD/count	Range
Circumference reduction rate (%)	30.0 \pm 5.1	18.0–39.7
Additional procedures after flap transfer		
Liposuction	1	
Contralateral breast surgery for symmetry	6	
Complications		
Seroma	0	
Infection	0	
Lymph node flap venous congestion	1	
Abdominal donor-site wound dehiscence	1	
Flap failure	0	

SD, standard deviation.

enhancements in microsurgical techniques. A simultaneous VLNT to treat lymphedema in patients who desire autologous MBR represents an enhancement in the comprehensive management of patients with breast cancer-related lymphedema (BCRL). This approach enables the achievement of both objectives in a single-stage procedure: breast reconstruction and concomitant treatment of lymphedema. Previously published studies with the use of the groin lymph node flap have shown this simultaneous surgical approach is feasible, safe and have promising results in patients' quality of life and surgical outcomes (21,22, 24–27).

The first study describing the use of abdominal-based free tissue transfer for breast reconstruction and simultaneous vascularized lymph node flaps was published in 2012 by Saaristo *et al.* (26). In this study, both DIEP and muscle sparing transverse rectus abdominis musculocutaneous (msTRAM) flaps were harvested along with lymph nodes surrounding the superficial circumflex iliac vessels (SCIV). The abdominal flaps were anastomosed to the thoracodorsal vessels and the lymph node flaps were anastomosed to the retrograde thoracodorsal vessels if needed. Seven of their nine patients showed a significant reduction in circumferential extremity measurements and five out of six patients with radiologic evaluation showed improved flow in lymphoscintigraphy at 6 months postoperative compared with preoperative pictures.

Since the introduction of this combined surgical approach, diverse lymph node flaps within which the most common are the inguinal lymph nodes along the SCIV or superficial inferior epigastric vessels (SIEV) have been used. One of the most important comparative advantages offered by these lymph node flaps is that the DIEP flap can be raised with the groin flap as a chimeric flap and transferred *en bloc*

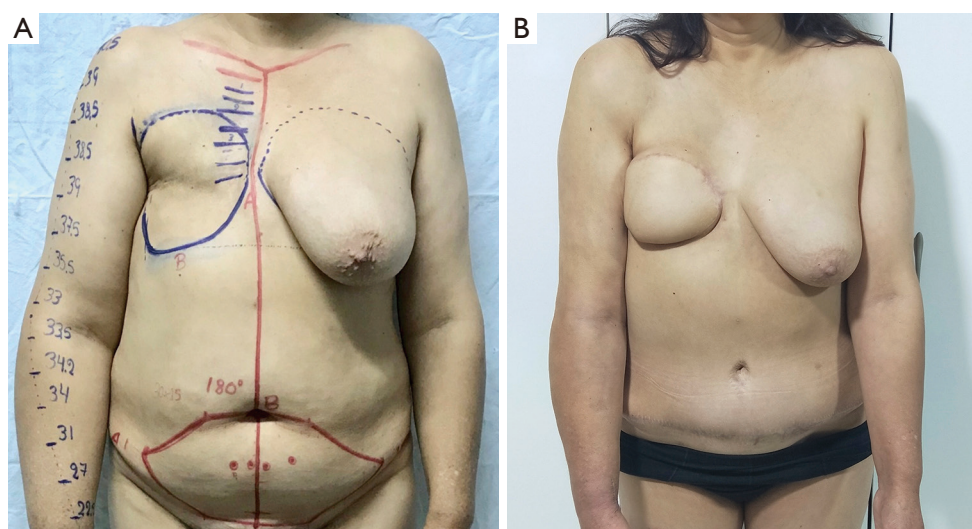


Figure 3 A 49-year-old female patient with left breast cancer-related lymphedema (BCRL) with 12 months of complex decongestive therapy (CDT) without significant improvement. The patient underwent the combined microvascular autologous breast reconstruction and gastroepiploic vascularized lymph node flap transfer. (A) Preoperative picture. Notice the mastectomy scar over her left chest and the volume and size discrepancy between the upper extremities. (B) Postoperative picture at 12-months follow-up evidencing a mean limb circumference reduction rate of 30%. In addition, the patient reported improved lymphedema-associated symptoms.

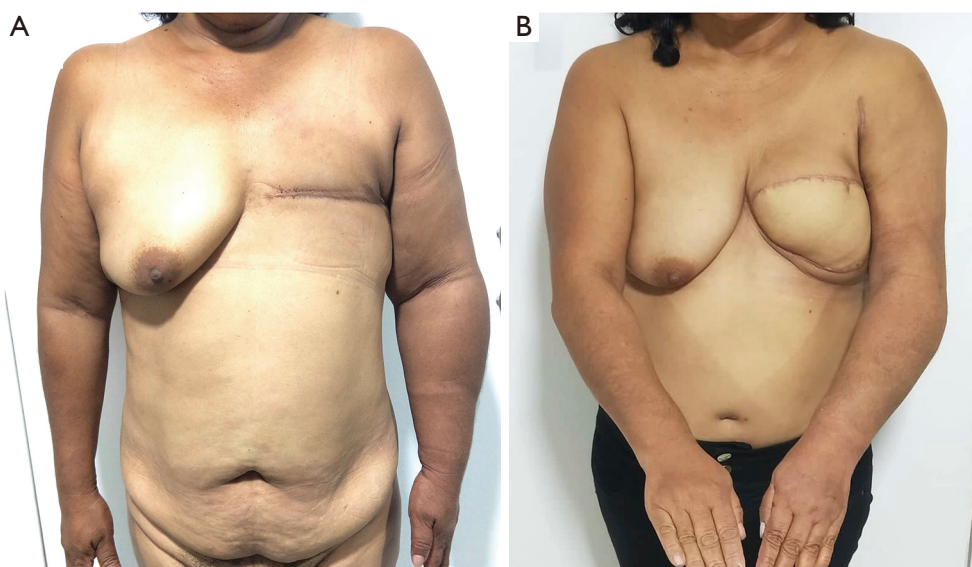


Figure 4 Pre and postoperative pictures of a 45-year-old female patient with right breast cancer-related lymphedema (BCRL). She had received 15 months of complex decongestive therapy with unsatisfactory results. (A) Preoperative picture. (B) Postoperative picture at 12 months of follow-up showing a limb circumference reduction rate of 39%.

to the recipient site. However, donor-site volume defect and risk of iatrogenic lymphedema are among its disadvantages. Careful dissection during flap harvest should be conducted to minimize the risk of iatrogenic lymphedema, which is

one of the most dreaded complications of VLNT. In order to avoid harvest of sentinel lymph nodes draining the lower extremity and decrease the risk of iatrogenic lymphedema, reverse lymphatic mapping can guide the harvest of

vascularized lymph node flaps. However, the risk of iatrogenic lymphedema is still latent (32). In addition, flaps based off of the SEIV have been reported to be associated with increased seroma formation (26,33).

Due to the increased risk of iatrogenic lower extremity lymphedema and the restrictions on shaping of the reconstructed breast with the use of chimeric abdominal flaps, the use of GE-VLNT during autologous breast reconstruction emerges as a strong surgical alternative. In our study, we used this flap showing a significant mean circumference reduction rate ($30.0\% \pm 5.1\%$, $P < 0.001$). To date, there have been no reports of iatrogenic lymphedema with the harvest of gastroepiploic vascularized lymph node flaps. It is versatile and contains a reliable amount of lymphatic tissue (at least three lymph nodes when 3 cm of surrounding tissue are obtained) (34). In addition, this flap can be divided into two isolated flaps based off of the two gastroepiploic vessels. These could be used as a double-inset in patients with unilateral extremity lymphedema (23) or even in patients with bilateral UEL who desire autologous bilateral breast reconstruction. In addition, the bulky appearance of the flap at the recipient site can be improved with microsurgical debulking if necessary (4). In our case series, we used a single-level distal inset as these sites are usually healthy, have no scar tissue and are usually spared from the deleterious effects of radiation therapy. Additionally, axillary scar release was done as it has been shown to be an effective strategy in order to fasten the onset of improvement and positively impact patient's quality of life (35).

Even though the harvest of the gastroepiploic vascularized lymph node flap is relatively unchallenging and offers the above-mentioned versatility, this procedure is more invasive and entails a mini laparotomy of approximately 5–7 cm. A detailed anatomical knowledge and a technically adequate abdominal wall closure are imperative to decrease donor site morbidity.

In our experience, for advanced lymphedema stages both the fluid and solid components of lymphedema should be addressed accordingly in order to obtain better outcomes. The VLNT adequately tackles fluid congestion and regenerates lymphatic connections; however, its role is still limited in cases where there is a high component of fibro-adipose deposition. In these cases, a combined approach including SAL or another debulking procedure may be required. One of our patients underwent SAL procedure 12 months postoperatively to improved surgical outcomes.

This study adds to the increasing body of evidence

supporting VLNT as an effective surgical treatment for lymphedema, particularly in patients undergoing breast reconstruction. However, it is not without limitations. The limited number of patients may not be representative and affects external validity. In addition, the follow-up time was relatively short. To the best of our knowledge, the present study is the largest case series with the longest follow-up combining microvascular breast and lymphatic reconstruction with DIEP flap and GE-VLNT.

Conclusions

The use of GE-VLNT for the treatment of lymphedema has shown promise in that there is no risk of iatrogenic lower extremity lymphedema. Based on our data, this is a safe and reliable procedure that adequately addresses two objectives: breast and lymphatic reconstruction. This combined surgical technique represents a great alternative to previously described donor-sites for the treatment of patients with BCRL who desire and are suitable for autologous MBR. This study paves the way for further prospective studies with longer follow-up to determine the true value of this combined surgical approach and to assess the differences between various donor flaps for VLNT in combination with MBR.

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Footnote

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Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The institutional review board approved the study (ALNH No: DMR005-IRB-103) and written informed consent was obtained from all patients.

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