Immediate Limb Compression Following Supermicrosurgical Lymphaticovenular Anastomosis – Is It Helpful or Harmful?

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Introduction
Lymphaticovenular anastomosis (LVA) is a minimally invasive, supermicrosurgical alternative to vascularized lymph node transfer (VLNT) for lymphedema reconstruction [1-3]. The procedure is conceptually simple, and it involves making lymph-to-vein connections via small skin incisions. In contrast to VLNT [4,5], the procedure does not involve harvesting lymphatic tissue and is therefore free of the risk of causing donor-site lymphedema. While its minimally invasive nature is appealing, its reported outcome is disappointingly inconsistent [6-8]. The inconsistency in outcome is likely related to the differences in surgical technique and peri-operative care. One of these controversies is whether to apply limb compression. Intuitively, limb compression following the LVA procedure may mechanically narrow these minuscule anastomoses, causing anastomotic failure. In standard microsurgery, pressure avoidance at the anastomotic site is an unchallenged dictum. But does the same hold true for the supermicrosurgical LVA? We conducted a simulation study to answer this question – does limb compression following LVA promote or impair lymph-to-vein drainage?

Method

Patients
Five consecutive patients, four female and one male, with age ranging from 23 years to 69 years; undergoing LVA for limb lymphedema, were included in the study (Table 1). All patients had previously failed complex decongestive lymphedema therapy and were referred by our lymphedema therapists for evaluation for surgical reconstruction. All had lower extremity lymphedema. Three had acquired disease and two had primary disease. The severity of disease was staged with Campisi criteria and all had stage II and III diseases.

Table 1. Comparison of Pre-operative and Post-operative LEL Indices and LYMQOL Scores

| Patient | Age | Extremity | Etiology     | Campisi Stage | No. of LVA | LEL Indices (Pre/Post) | LYMQOL (Pre/Post) |
|---------|-----|-----------|--------------|---------------|------------|------------------------|------------------|
| 1       | 52  | Leg       | Acquired     | II            | 10         | 294/279                | 82/57            |
| 2       | 69  | Leg       | Acquired     | II            | 8          | 312/294                | 97/62            |
| 3       | 23  | Leg       | Congenital   | III           | 9          | 337/312                | 103/79           |
| 4       | 58  | Leg       | Congenital   | II            | 7          | 298/282                | 78/46            |
| 5       | 67  | Leg       | Acquired     | III           | 8          | 319/305                | 94/53            |
| P       |     |           |              |               |            | 0.0009                 | 0.0006           |

Lymphedema index is a circumference-based system that takes measurements at five limb levels and references the sum to the patient’s body mass index. LYMQOL is a validated lymphedema-specific quality of life assessment that tracks four condition-specific domains – function, appearance, symptoms, and mood. LEL, lower extremity lymphedema; LYMQOL, lymphedema-specific quality of life assessment; LVA, lymphaticovenular anastomosis; Pre, pre-operative; Post, post-operative.
Study Design

Intraoperatively, immediately following the completion of each of the individual LVAs, the flow pattern across the anastomosis was observed under the surgical microscope. When the lymphatic pressure exceeded the venous pressure, favorable antegrade lymph-to-vein flow occurred and a “washout” sign (Figure 1A) was observed. Conversely, when the pressure gradient was reversed, with the venous pressure exceeding the lymphatic pressure, unfavorable retrograde flow occurred and a “backflow” sign was seen (Figure 1B). Regardless of the initial flow pattern observed, bandage compression was applied to the entire limb to simulate postoperative limb compression (Figure 2). The firmness of compression was determined by the senior author to simulate the compression pressure of 30 – 40 mmHg. Changes to the flow pattern following the bandage compression were observed under the microscope and recorded. Patient evaluation was performed using the circumference-based lower extremity lymphedema (LEL) index system [9] and a lymphedema-specific quality of life assessment (LYMQOL) [10] at preoperative visit; and at the 3rd and 6th month postoperative visits. LYMQOL is a condition-specific validated assessment system that tracks postoperative changes in function, appearance, symptoms, and mood. The preoperative and 6-month postoperative LEL and values were compared using paired t-test.

Surgical Technique

After mapping the lymphatic vessels with indocyanine green lymphography and delineating the superficial venules with an infrared imaging device (VueTek Scientific, Gray, Maine), the incisions were strategically placed at locations where both the lymphatic vessels and venules were present as previously described [8] (Figure 3). 0.05 cc of isosulfan blue (Lymphazurin; United States Surgical Corp., Norwalk, Connecticut) was injected within 2 cm distal to each incision to further facilitate identification of the lymphatic vessels. The LVAs were performed at 25X magnification utilizing a surgical microscope (Pentero 900; Carl Zeiss, Oberkochen, Germany) using specialized supermicrosurgical instruments (EMI Factory, Kitasakugun, Nagano, Japan). Both the standard supermicrosurgical LVA technique described by Koshima and Yamamoto et al. [11-13] and the “octopus” technique [14] were used. When healthy lymphatic vessels and size-matched veins were present, the standard technique was pref-
of limb compression, 16 of the 17 LVAs (94%) converted the flow pattern the "octopus" and 4 were the standard LVAs. Following the application

sign. Among these 17 LVAs demonstrating the "backflow" sign, 13 were and the remaining 17 (40%) demonstrated the unfavorable "backflow"

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Postoperative Care

Limb compression was applied immediately following the surgery using the short-stretch bandage. All patients were discharged to home on postoperative day one. Bandage compression continued for 16 hours per day until six weeks postoperatively. At that time, all were transitioned to 30-40 mmHg pressure. Throughout the 6-month study period, all patients continued to wear their pressure garments for 16 hours per day. Wean-

ing "washout" was seen in all 3 of the lymphatic vessels. Note how the blood was "washed out" of all 3 of the lymphatic vessels and the vein. Engagement of the lymphatic vessels was clearly seen in the top 2 lymphatic vessels.

Results

A total of 42 LVAs were created – 26 standard LVAs and 16 "octopus" LVAs. Initially, 25 of the LVAs (60%) demonstrated the favorable "washout" sign and the remaining 17 (40%) demonstrated the unfavorable "backflow" sign. Among these 17 LVAs demonstrating the "backflow" sign, 13 were the "octopus" and 4 were the standard LVAs. Following the application of limb compression, 16 of the 17 LVAs (94%) converted the flow pattern from the unfavorable "backflow" to the favorable "washout" (Figures 4A & 4B). The conversion rates of the "octopus" and the standard LVAs were 100% (13/13) and 75% (3/4), respectively. All 25 LVAs initially demonstrating "washout" maintained the "washout" pattern following the bandage compression. Regardless of the initial flow patterns, post-compression vessel engorgement was observed in the lymphatic vessels (Figure 4B). This finding along with the post-compression flow reversal suggested a compression-induced augmentation of the lymphatic pressure.

All patients demonstrated reduction of limb swelling and relief of symptoms, as demonstrated by statistically significant improvements in the lower extremity lymphedema indices (P = 0.0009) and in the LYMQOL (P = 0.0006) (Table 1). All patients reported their symptoms being notably more responsive to compression. No patient had worsening of lymphedema symptoms during the 6-month study period.

Discussion

LVA is a delicate supermicrosurgical procedure. Until now, most studies focused on the intricate technical aspects of the procedure [13-16] and little had been described about postoperative management. As we become more proficient in creating these tiny anastomoses, it is important to start to evaluate other procedural parameters to maximize surgical efficacy. The opinions and practices on postoperative limb compression following LVA vary widely among the supermicrosurgeons. Common practices include no compression [17,18], delayed compression starting few weeks following the surgery [11], and immediate compression [6]. To our knowledge, this is the first simulation study directly evaluating the effects of compression on the LVA.

The findings in this study support immediate postoperative limb compression. When the compression was applied, majority of the unfavorable LVAs with retrograde flow converted to the favorable functioning LVAs with ante grade flow (94%). When left untreated, the refluxed blood in the lymphatic lumen may cause thrombosis and result in anastomotic failure. Interestingly, the limb compression appeared to create a general-

ized augmentation of the lymph-to-vein pressure gradient, as suggested by visible engorgement of lymphatic vessels (Figure 4B). This phenomenon was seen in all the LVAs, including those already demonstrating ante grade flows prior to compression. This finding along with the high rate of favorable flow conversion suggested functional enhancement of the LVAs with postoperative limb compression.

Why would limb compression, which pressurized both the venous and lymphatic systems simultaneously, alter the lymph-vein pressure situation?
gradient? We hypothesized that the differential effects on the two systems in response to external pressure may be related to the lymphatic system being a partially obstructed system and the venous system being a non-obstructed, free-flowing system. When both systems were simultaneously pressurized, the partially obstructed system experienced a higher magnitude of pressure increase relative to the open system due to its inability to efficiently decompress (Hai Fu, Department of Physics and Astronomy, University of Iowa, personal communication, December 22, 2016) (Figure 5). In contrast, not having an outflow obstruction, the open venous system could quickly decompress and therefore experienced a lesser pressure increase.

We were not surprised by the higher incidence of “backflow” in the “octopus” LVA (81% or 13 of 16) relative to the standard LVA because the lymphatic vessels used in the “octopus” technique were qualitatively worse than the ones used in the standard technique, and were mostly of the “contraction type” [19,20]. Without compression, these LVAs would likely not be effective due to their inability to peristaltize and generate a favorable lymph-to-vein pressure gradient. Using compression, we were able to convert 100% (13 of 13) of the unfavorable “octopus” LVAs with “backflow” to the favorable, functioning LVA showing “washout”. This finding is encouraging because it suggested that even the damaged “contraction type” lymphatic vessels may be successfully recruited to build functioning LVAs. This allows less restrictive lymphatic vessel recruitment and will lead to increased number of functioning anastomoses, and is a recommended postoperative practice.

In summary, the benefits of immediate compression following the LVA are three-fold:1) it converts nonfunctioning LVAs with retrograde flow to functioning ones with ante grade flow, 2) it augments the flow of the functioning LVAs already demonstrating ante grade flow, and 3) it decreases the restrictive nature of the LVA procedure and allows the surgeon to use moderately disease-affected lymphatic vessels, creating higher numbers of functioning LVAs. Currently, the endpoint of limb compression and the timing of its discontinuation are unknown, and they are being investigated in our ongoing studies.

**Conclusion**

Immediate limb compression following the LVA procedure facilitates lymphatic drainage and increases the surgical efficacy by increasing the number of functioning anastomoses, and is a recommended postoperative practice.

**Article Information**

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Lymphedema; lymphaticovenular anastomosis; lymphedema surgery; supermicrosurgery.
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