#### ORIGINAL RESEARCH

# Minimally Invasive Treatment of Chyle Leak After Thyroidectomy and Cervical Lymph Node Dissection in Patients with Thyroid Carcinoma: Results of a Study Involving 36 Patients

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**Objective:** Chyle leak (CL) after head and neck surgery is a rare but well-known complication. In patients with high-output leakage, the treatment can be complicated. This study aims to report on a recent innovation in lymphatic intervention for treating such patients. **Materials and Methods:** A retrospective review of 36 patients with chyle leak after neck surgery for thyroid cancer was conducted to assess the efficacy of percutaneous lymphatic embolization and thoracic duct (TD) disruption.

**Results:** Antegrade catheterization of the thoracic duct was achieved in 31 of 36 patients (86.1%). Therefore, embolization of the thoracic duct branches was performed in 26 and 5 patients, respectively. In 5 cases of unsuccessful antegrade catheterization into the thoracic duct, transcervical access embolization was performed in 2 patients, and TD disruption (TDD) was performed in 3 patients. The pooled overall technical success rate of lymphatic embolization was 33/36 patients (91.7%). One patient who underwent thoracic duct embolization (TDE) with technical success (1/33 patients) but clinical failure had additional treatment directly sclerosing the TD under computed tomography scan. Cervical fluid collection sclerotherapy was done in 7 patients as an additional treatment. Resolution of the chyle leak after procedures was observed in all patients (100%). The mean time to resolution was 3 days (1–7 days). There was no complication intra and after procedures.

**Conclusion:** TDE, selective TD branches embolization and TDD are safe and effective minimally invasive treatments for CL postsurgery for thyroid carcinoma. Sclerosing cervical fluid collection contributes to clinical success.

Keywords: thyroid cancer, thyroidectomy, chyle leak, thoracic duct embolization, thoracic duct disruption, sclerotherapy

#### Introduction

Chyle leak is a rare but serious complication of thyroidectomy. It is reported that the incidence is 0.5–1.4% of thyroidectomies and the risk increases when thyroidectomy is associated with neck dissection, up to 8.3%.<sup>1–3</sup> Chyle leak may be identified intraoperatively, postoperatively immediately or several days after surgery when enteral feeding is started.<sup>4</sup> Prolonged high-volume loss of chyle leads to nutritional failure and immunosuppression, which may even be life-threatening. The output volumes and color of the drainage depend on diet and injury site of the thoracic duct or its branches.

Currently, there are no guidelines for management of chyle leak post-thyroidectomy. The treatment options for postthyroidectomy chyle leak depend on drain output and patient comorbidities. Management usually starts with conservative measures. Conservative treatments such as nutritional management, pressure dressings, negative pressure suction dressing,

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somatostatin and octreotide may be effective in patients with low output.<sup>4</sup> Patients who had persistent high output or failure of conservative measures required other management approaches such as surgery, thoracic duct embolization, or sclerotherapy.<sup>5,6</sup> Lymphatic embolization has been reported as an effective method of managing patients with chylous ascites and chylothorax.<sup>7,8</sup> There are some sporadic reports about TDE in management of CL post-thyroidectomy when conservative management is ineffective.<sup>9,10</sup> The purpose of this study was to evaluate the technical success and clinical outcomes of minimal invasive intervention in management of CL following cervical lymph node dissection in thyroid carcinoma patients.

## **Materials and Methods**

#### **Patient Characteristics**

The retrospective study included 36 patients with chyle leak after thyroidectomy and cervical lymph node dissection in patients with thyroid carcinoma. This was a review of all patients who underwent TDE or TDD for the management of CL at Hanoi Medical University Hospital from June 2019 to July 2023. All subjects had confirmed CL based on drain fluid triglyceride levels greater than 110mg/dL. Conservative therapy has failed to stop chylous leakage in these patients, which included a low-fat diet or total parenteral nutrition and octreotide, and reoperation. Our retrospective study was in accordance with the ethical guidelines of the Declaration of Helsinki. This retrospective study was approved by the institutional review board of the Department of Science and Technology of Hanoi and Hanoi Medical University Hospital (Ref: 139/QĐ-SKHCN dated 24 February 2023) and all patients provided written informed consent.

Data was obtained from patient records including age, sex, surgical procedure, clinical symptoms, volume of drainage output (high-output  $\geq$ 500 mL/day, low-output <500 mL/day), lymphangiography findings, variant anatomy of thoracic duct, TDE/TDD technical approach, material used to perform the procedure, subsequent interventions, complications, and hospital stay. The amount and duration of chyle drainage output volume per 24 hours before and after the procedure were recorded.

## **Techniques**

Thoracic duct embolization (TDE) and TDD were used.

### Transabdominal Antegrade Access

Ultrasound of the bilateral inguinal lymph node was performed to identify lymph nodes. Then, a needle accessed the inguinal node under ultrasound guidance. A total of 10 mL oiled contrast agent was slowly injected into the bilateral inguinal lymph node under fluoroscopic guidance. After the cisterna chyli and the thoracic duct became visible, a 21-gauge Chiba accessed the cisterna chyli via the transabdominal approach. A guidewire was advanced through the needle into the cisterna chyli. Then, a microcatheter was advanced over guide wire into the TD. Contrast was injected through the microcatheter to confirm the site of leakage and evaluate the anatomy of the thoracic duct. TDE was then performed using different embolization materials.

## Transcervical Retrograde Access

Transcervical access was performed after a failed antegrade access attempt. A 21-gauge Chiba was directly punctured to the distal of the TD near its insertion at the venous angle under fluoroscopic guidance. Then, a microcatheter was advanced over a guide wire into the TD. Lymphatic embolization was then performed.

## Thoracic Duct Disruption (TDD)

TDD was performed as described by Cope et al.<sup>11</sup> In patients whose thoracic duct cannulation was unsuccessful, TDD was performed by puncturing the TD, the tissue next to the TD multiple times with a fine needle, and injection of a mixture of Histoacryl and Lipiodol at a 1:3 ratio or saline in the space surrounding the TD under fluoroscopy. This may reduce or interrupt the lymphatic flow by leaking the sclerosing agents into the TD via scratching or compression of the thoracic duct, allowing the leak to heal spontaneously.

### Outcome

The technical success of lymphatic embolization was defined as cannulation of the TD via transabdominal antegrade or cervical retrograde access and delivery of embolic material, if indicated, to treat a chyle leak. The clinical success of the procedure was defined as the drain output being resolved within 1 week after procedures.

## Statistical Analyses

Demographic and operative data, location of the chyle leaks, drain output data, management approach, embolization materials, technical success (the ability to access the TD and subsequent embolization of the TD), and clinical success (resolution of the presenting chyle leak) were retrospectively collected. These data were analyzed using standard statistical techniques, including medians and ranges for continuous data and counts and percentages for categorical data. Follow-up data were obtained until patients' discharge and at re-examination one month later.

## Results

The study included 36 patients (30 females and 6 males), with a mean age of 49.8 years (range: 21 to 77 years). Of the total 36 patients, 31 had experienced a thyroidectomy with cervical node dissection and 5 patients had undergone only cervical node dissection, with no history of thyroidectomy. Most patients (29) presented with creamy or milky drain output, 7 patients presented with neck palpable fluid collection, and 3 patients had dyspnea. Most patients developed a leak on the left side (34 patients, 94.4%) and 2 patients had left cervical and thoracic leaks. All patients were switched to a fasting or low-fat diet. Four patients experienced repeat surgery for ligated or stitched thoracic duct; one patient was reoperated on twice. Patient demographics and clinical presentations are summarized in Table 1.

The maximum drain volumes ranged from 100 to 1500 mL per 24 hours (mean  $342 \pm 338$  mL), and the duration of leakage prior to intervention ranged from 7 to 52 days (mean  $18.5 \pm 11$  days). The duration of leakage after intervention ranged from 1 to 7 days (mean  $3 \pm 1.7$  days). After the procedure, the output drainage decreased (mean  $20 \pm 19$  mL; range 10-75 mL). The characteristics of chyle leak are summarized in Table 2.

On lymphangiography, most patients had an injury of the terminal thoracic duct (27 patients, 75%) and 9 patients had CL due to injury of a thoracic duct branch. Six patients had an anatomical variance: the cisterna chyli was not visualized in 3 patients, 2 patients had a plexiform thoracic duct in the mediastinum, and 1 patient had a double thoracic duct with bilateral terminations to the left and right subclavian veins.

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Characteristics		Value
Age (years)		49.8 ± 12.6 (21–77)
Gender		30 female, 6 male
Original surgery	Thyroidectomy + neck dissection	31
	Neck dissection only	5
Presentation	Cervical fluid collection	7
	Creamy or milky drain output	29
	Dyspnea	3
Presentation site (n)	Left neck	34
	Right neck	I
	Thoracic	3
Management before intervention (n)	Conservative	32
	Surgery	4

 Table I Characteristics of Patients with Chyle Leak

Characteristics		Mean ± SD (range)	
Type of leakage	High output	II patients (mean 900 ± 516 mL)	
	Low output	25 patients (mean 186 ± 93 mL)	
Maximum drain volume prior intervention (mL per day)		342 ± 338 (100–1500)	
Duration of leakage prior to intervention (days)		18.5 ± 11 (7–52)	
Duration of leakage after intervention (days)		3 ± 1.7 (1–7)	
Maximum drain volume after intervention (mL)		20 ± 19 (10–75)	

Table 2 Characteristics of Chyle Leak

Catheterization of the TD was technically successful in 31 patients (31/36). Transabdominal embolization was performed in these patients: 26 underwent thoracic duct embolization and 5 experienced thoracic duct branch embolization. The reasons for failed TD access were small or absent cisterna chyli. Five patients experienced failed TDE: two of them underwent transcervical TD access (1 patient with TD branch embolization and 1 patient with TDE) and 3 others had TDD. For patients with neck fluid collection, an additional sclerotherapy treatment for fluid collection with absolute ethanol was performed (in total, 7 patients: 6 with TDE and 1 with TDD). The materials used in the embolization procedures were n-BCA glue in 14 patients, and coils combined with n-BCA glue in 19 patients. The characteristics of the procedures are summarized in Table 3.

The combined technical and clinical outcomes of intervention are summarized in Figure 1. The technical success rate of transabdominal embolization was 86.1% (31/36 patients); both transcervical access (2/2) and TDD (3/3) were 100% successful. The overall technical success rate of lymphatic embolization was 91.7% (33/36). Following transabdominal embolization, 30 of 31 patients experienced resolution and 1 patient did not. This patient underwent a second procedure: sclerosing injection to the thoracic duct under computed tomography guidance on the 23rd day after TDE. Then, this patient received total parenteral

Characteristics		N (patients)
Findings on lymphangiography	Injury of the thoracic duct is terminated	27
	Injury of the thoracic duct branch persists	9
Anatomical variation	Absent cisterna chyli	3
	Plexiform presentation	2
	Double thoracic ducts with bilateral termination	I
Management of intervention	Embolization of TD	26
	Embolization of TD branches	5
	Transcervical access embolization	2
	TDD	3
Embolization materials	NBCA	14
	NBCA+ coils	19
Additional management	Sclerosing injection of collected fluid	7
	Sclerosing injection of TD	I

Table 3 Procedure Characteristics

Abbreviations: TD, Thoracic duct; TDD, Thoracic duct disruption; NBCA, N-butyl cyanoacrylate.

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Figure I Flowchart of the technical and clinical outcomes of 36 patients. One patient underwent 2 procedures.

nutrition and octreotide. The causes of failure were inadequate filling and washout of the glue in the lumen of the thoracic duct when the patient was discharged and returned to a normal diet. There was no complication intra and after all procedures.

The clinical success rate after successful transabdominal retrograde access lymphatic embolization was 30/31 patients (96.8%), and after TDD was 3/3 patients (100%). Two patients experienced successful resolution of CL after undergoing transcervical embolization. Clinical success after the initial procedure was found in 35/36 patients. In one patient who underwent a second procedure after initial clinical failure, clinical success was also achieved. Overall, clinical success was achieved in 100% of patients. The hospital stay after intervention was from 3–15 days (median 5 days). Follow-up was performed after 1 month of discharge no one has experienced persistent or recurrent CL.

#### Discussion

Chyle leak develops as a result of injury to the lymphatic duct after head and neck surgery. It may present as cervical or thoracic leaks. Incidence is significantly higher on the lefthand side because of the anatomy of the TD,<sup>10,12</sup> as was the case in our study. One patient presented with a chyle leak on the right side of the neck because of thoracic duct anatomical variance (double thoracic duct with bilateral termination). In our series, we found various leakage points resulting from differences in thoracic duct anatomy (Figure 2). There is no consensus on the need for reoperation or lymphatic embolization as defined by chyle volume loss or timing. Indications for non-conservative treatment vary widely among studies, ranging from 100 mL to 1.1 L per day and/or a persistent chyle leak after 2 to 14 days of conservative management.<sup>13,14</sup> Reisenauer et al<sup>15</sup> recommended that chyle drainage of  $\geq$ 1100 mL in 24 hours postoperatively should be considered the threshold for



Figure 2 Typical features of thoracic duct lesion. (A) Laceration of the proximal part of the TD (arrow) causing extravasation of contrast in fluid collection (star). (B) Laceration of the TD branch (long arrow) causing extravasation of contrast (star) while the normal terminal part of the TD was intact (short arrow). (C) Anatomic change of the TD with the branches; one of these branches was ruptured after lymph node resection (arrow). (D) TD occlusion (short arrow) and laceration of collateral lymphatic vessels (long arrow).

intervention. Others suggest that lymphatic embolization is indicated when the drainage output is >1000 mL in 24 h or >100–300 mL/24 h for over 1 week.<sup>14,16</sup>

In our study, transabdominal access was performed in all patients but success was achieved in 86.1% (31/36). Transcervical access and TDD were performed and successful in 2 and 3 patients, respectively. The overall technical success rate of lymphatic embolization was 91.7% (33/36). This rate is lower than that reported by Nadolski et al<sup>17</sup> (98%) but higher than the 62.9% reported following the meta-analysis conducted by Kim et al,<sup>18</sup> the 67% found by Itkin et al,<sup>19</sup> and the 79% reported by Pamarthi et al.<sup>20</sup> Crawford et al<sup>21</sup> reported an overall TDE success rate of 68%; TDE with transabdominal antegrade access had a 44% success rate; and TDE with transcervical retrograde access had a 77% success rate. There are very few reports about using lymphatic embolization or TDD in management of chyle leak from neck surgery; almost all are sporadic case reports or case series.<sup>10,14</sup> Moussa et al<sup>10</sup> reported 6 cases undergoing a total of seven TDE procedures; resolution was achieved in all patients. The technical failures of transabdominal access in this study were due to unsuccessful cannulation of the cisterna chyli or thoracic duct. The reasons were small cisterna chyli or inability to visualize them or anatomic variations of the TD (eg, plexiform presentation, double TD). The study of Crawford et al<sup>21</sup> showed that most patients who failed TD access had small cisterna chyli; other reasons were TD occlusion and slow progression of the contrast agent to the cisterna chyli. Pamarthi et al<sup>20</sup> reported that unsuccessful cannulation was caused by no visualization of the cisterna chyli, inadequate pedal lymphatic vessels for cannulation, left paraaortic cisterna, oversedation, and power injector

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malfunction. In addition to TDE, we tried to access and embolize the injury branch of TD in order to preserve the thoracic duct; technical and clinical success was achieved in 6 patients. This method has not been commonly mentioned in the literature describing treatment of chyle leak. Kim et al<sup>14</sup> reported a study of 8 patients who were successfully treated with lymphatic embolization to manage chyle leak after neck surgery; selective embolization of the TD branches (5 patients) was more highly prioritized than embolization of the terminal thoracic duct (2 patients). Selective embolization with preservation of the thoracic duct has potential because it maximizes preservation of the lymph flow. Moreover, percutaneous ethanol injection of the collected cervical fluid causes fibrosis of the connective tissue around the fistula, which contributes to healing the leak.

The overall clinical success rate of procedures in treatment of chyle leak in this study was 100% for all subjects. This finding is higher than those of Pamarthi et al (64%),<sup>20</sup> Itkin et al  $(71\%)^{19}$  and Yannes et al (71%).<sup>22</sup> A meta-analysis conducted by Kim et al<sup>18</sup> showed that the clinical success rates of TDE, TDD and overall were 79.5%, 60.8% and 60.1%, respectively. A systematic review showed median clinical success rates for TDE of 74.6% (range: 57–98%) and TDD of 72% (range: 41.7–100%).<sup>13</sup> The use of multiple techniques as assistance or alternative methods when transabdominal access fails could reduce the clinical failure rate. According to Kim et al<sup>14</sup> and Moussa et al,<sup>10</sup> chyle leak resolved in all patients from 1 to 5 days after intervention. Another study demonstrated that the median time to resolution of chyle leak in patients with TDE was 3 days, and in patients with TDD was 7 days.<sup>22</sup>

There were no minor or major complications in our study. The rate of complications related to TDE and TDD ranged from 0% to 6.7% and most were minor complications.<sup>10,14,19,20</sup> These included non-target glue embolization to the lung, non-target glue embolization to the portal vein, retained guide wire fragment in the retroperitoneum, transient apnea caused by oversedation, perihepatic hematoma at the percutaneous access site, bile leak, further chyle leak, leg edema, and inconsequential coil misplacement.<sup>13,14,20</sup> Other possible complications of TDD reported are a hematoma between the aorta and vertebral body and a guide wire fracture.<sup>22</sup>

There are some limitations to this study. First, we used a retrospective design and there is a lack of long-term follow-up. The patients were followed for only 1 month after the procedure, so no data on long-term complications of TD embolization were collected. Second, the sample size was relatively small and involved a single center; it may thus not be representative of other centers. Third, most of the patients came from other hospitals and had received treatment over a long period of time; thus the previous approaches and surgical ligation techniques experienced by them are unknown and varied. Despite these limitations, this study may provide useful information for the management and treatment of chyle leak following thyroidectomy.

#### Conclusion

Lymphatic embolization and TDD is a safe and effective procedure that results in resolution of chyle leak after surgery for thyroid cancer. Sclerosing therapy is an adjuvant therapy that contributes to reducing the clinical failure rate. Lymphatic intervention should be the initial treatment of choice for high-output chyle leak or failure of conservative management. A long-term study should be conducted to assess the impact of thoracic duct embolization.

#### **Data Sharing Statement**

The datasets generated or analyzed during this study are available from the corresponding author on reasonable request.

## **Author Contributions**

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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# Disclosure

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