

Effects of a physical therapy program combined with manual lymphatic drainage on shoulder function, quality of life, lymphedema incidence, and pain in breast cancer patients with axillary web syndrome following axillary dissection

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Abstract

Purpose The aim of this study was to evaluate the effects of physical therapy (PT) combined with manual lymphatic drainage (MLD) on shoulder function, pain, lymphedema, visible cords, and quality of life (QOL) in breast cancer patients with axillary web syndrome (AWS).

Methods In this prospective, randomized trial, 41 breast cancer patients with visible and palpable cords on the arm and axilla and a numeric rating scale (NRS) pain score of >3 were randomly assigned to PT (3 times/week for 4 weeks; $n=20$) and PT combined with MLD (5 times/week for 4 weeks; PTMLD; $n=21$) groups. MLD was performed by a physical therapist and the patients themselves during week 1 and weeks 2–4, respectively. Arm volume, shoulder function (muscular strength; active range of motion; and disabilities of the arm,

shoulder, and hand [DASH]); QOL (European Organization for Research and Treatment of Cancer Core and Breast Cancer-Specific QOL questionnaires), and pain (NRS) were assessed at baseline and after 4 weeks of treatment.

Results QOL including functional and symptom aspects, shoulder flexor strength, DASH, and NRS scores were significantly improved in both groups after the 4-week intervention ($P<0.05$). NRS score and arm volume were significantly lower in the PTMLD group than in the PT group ($P<0.05$). Lymphedema was observed in the PT ($n=6$), but not PTMLD, group ($P<0.05$).

Conclusions PT improves shoulder function, pain, and QOL in breast cancer patients with AWS and combined with MLD decreases arm lymphedema.

Keywords Axillary web syndrome · Breast neoplasms · Physical therapy modalities · Quality of life

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Introduction

Axillary web syndrome (AWS) is a common postoperative complication in breast cancer patients. AWS usually occurs 5–8 weeks after surgery and is characterized by visible or palpable cords of subcutaneous tissue in the breast, medial arm, antecubital space, forearm, hand, or chest wall. This syndrome often limits shoulder and elbow range of motion (ROM), causing pain and tightness [1–4]. Previous studies have demonstrated that AWS incidence varies according to the type of axillary surgery (sentinel node lymph node dissection [SLND] or axillary lymph node dissection [ALND]). AWS incidence rates of 20 and 38–72 % have been reported following SLND and ALND in breast cancer patients, respectively [2, 5, 6]. Bergmann et al. [7] reported that the incidence of AWS after breast cancer surgery was 28.1 % and was related to ALND and numbness in the arm after intercostobrachial nerve injury. The pathophysiology of

AWS has not been extensively investigated, with only a few studies reporting AWS as a benign occurrence. AWS is reported to be associated with lymphovenous damage, lymphatic stasis, and tissue injury resulting from the disruption of superficial lymphatics during axillary surgery [8]. A recent study asserted that AWS and the palpable cords were associated with lymphatic origin, and that they are not related with superficial vein thrombosis or with fascial problem as in Mondor's disease [1]. Therefore, further studies of the etiology and pathophysiology of AWS are needed [9].

Currently, there is no definitive treatment for AWS. Nonsteroidal anti-inflammatory drugs (NSAIDs) and physiotherapy including ROM exercises were found not to be effective in relieving early symptoms of AWS in breast cancer patients [2, 10]. However, other studies have suggested the importance of a physical therapy-directed approach in the AWS management [11, 12]. Most breast cancer patients experience acute postoperative pain with minimal daily activity [13]. Postoperative pain can cause anxiety in using the affected arm, which can limit arm movement and consequently result in muscle shortening and decreased muscle activity [14]. Furthermore, acute postoperative pain can become chronic if left untreated, leading to persistent arm and shoulder disability [15]. Some studies have shown that physical therapy can shorten the natural course of AWS by up to 6 to 8 weeks [2, 6, 10]. In addition to physical therapy, manual lymphatic drainage (MLD) may also reduce the development of AWS [16]. Increased clearance is thought to reduce local levels of inflammatory mediators, which are often associated with edema and pain [17]. MLD has been shown to improve blood circulation [18], stimulate the movement of lymphatic and other tissue fluids [18], and promote fluid clearance and tissue softening [19] in patients with athletic injuries. Early stimulation of lymphatic drainage can also modulate pain and inflexibility associated with lymphatic vessel inflammation secondary to surgical injuries [20]. The therapeutic effects of MLD are likely due to the prevention of protein stagnation and inhibition of factors capable of producing lymphatic overload. The effect of MLD in breast cancer patients with AWS remains unknown. The purpose of this study was to compare shoulder function, lymphedema incidence, presence of visible cords, and quality of life between breast cancer patients with AWS receiving physical therapy alone or in combination with MLD. The purpose of this study was to evaluate the effects of physical therapy (PT) combined with MLD on shoulder function, pain, and lymphedema symptoms in comparison to PT only, in breast cancer patients with AWS.

Methods

Study design and patients

This prospective, randomized, controlled trial was conducted at the Department of Rehabilitation Medicine of Asan

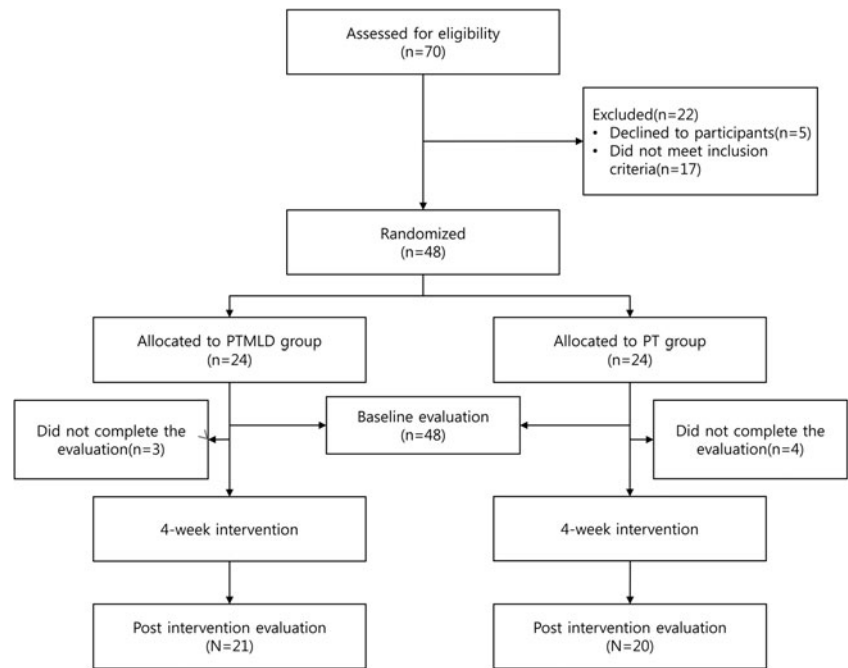
Medical Center, Seoul, Korea. Patients who visited the rehabilitation medicine department for regular check-up after surgery were recruited. Breast cancer patients with pain over 3 points using the pain rating scale of numeric rating scale (NRS) on upper/lower arm, elbow, and dorsum site in the region of the cording, and visible or palpable cords in the arm or breast at least 4 weeks after breast cancer surgery were included in the analysis. NSAIDs (two times daily) were prescribed for all patients. Patients with lymphedema, acute thrombosis, skin problems such as infection, or musculoskeletal disorders such as low back pain, disk pain, osteoarthritis, rheumatoid arthritis, pectoral muscle tightness, rotator cuff disease, and adhesive capsulitis were excluded. Also, cording only on the chest or lateral thorax and not involving an arm were excluded. A priori power analysis with G*Power ver. 3.1.5. software (Franz Faul, University of Kiel, Germany) was performed to calculate the sample size. The data of a pilot study of eight subjects was used to achieve the effect size of 1.1, the alpha level of 5 %, and the power of 80 %. The calculated sample was 30. Seventy patients were initially enrolled, of which 22 dropped out of the study because of the long hospital commute. The remaining 48 patients were randomized into two intervention groups: PT and PT combined with MLD (PTMLD). Seven patients were unable to complete the final evaluation; therefore, only 41 patients completed the study (Fig. 1). All measurements were taken by one therapist who was blinded. Written consent was obtained from all patients, and ethics approval of the study was obtained from the Asan Medical Center (2014-0551).

Interventions

Physical therapy

Both the PT and PTMLD groups underwent a physical therapy program three times a week for 4 weeks. Physical therapy sessions included a 10-min warm-up and cool-down, which consisted of eight stretching exercises (Online Resource 1), and strengthening exercises. For upper extremity strengthening, participants performed three different pulley exercises 10 times for three sets. For shoulder flexor, shoulder abductor, and elbow flexor strengthening exercises, participants performed three sets of 10 repetitions using a Thera-Band (Hygenic Corporation, Akron, OH.) Intensity of exercise was 6–8 on the OMNI Resistance which is equivalent to exercise intensity levels ranging from 60 to 80 % of 1RM (24) for active muscle scale, indicating 'somewhat hard'. OMNI Resistance of 0 indicates 'extremely easy' and 10 indicates 'extremely hard'. All exercises were performed under the supervision of a physical therapist. Manual therapy was then performed for 30 min by a skilled physical therapist and involved gentle circular mobilization of identified tight and stiff tissues of the chest wall and antecubital fossa with full hand or

Fig. 1 Flow of participants throughout the trial. *PTMLD* physical therapy combined with manual lymphatic drainage, *PT* physical therapy



two finger contact and longitudinal tissue stretch to the tight cords with the patient's arm in possible range of abduction [19]. Treatment intensity was progressively increased from comfortable to mild discomfort within the tolerable range. In addition, scapular mobilization was used to decrease scapular stiffness. Facing the patient in a side-lying position, the physical therapist pushed underneath the medial border of the scapula and released with the fingertips. The manual therapy program for AWS was performed in the following order: (1) soft tissue mobilization techniques and stretching for tight tissue cords; (2) shoulder abduction, elbow extension, and wrist supination and extension stretching exercises; (3) shoulder girdle mobilization; and (4) passive ROM (PROM) exercises.

Manual lymphatic drainage

MLD was provided by a certified lymphedema therapist. The PTMLD group additionally received 30 min of MLD daily five times a week for 4 weeks. MLD is a light form of circular massage (superficial tissue stretching) and involves stationary circle, pumping and scooping, and rotary movements performed with varying degrees of pressure. MLD was performed by two certified physical therapists (Dr. Vodder Method) during week 1. For weeks 2–4, the patients themselves performed MLD.

Assessments

Arm volume, muscular strength, active ROM (AROM), pain, visible cording, arm disability, and quality of life were evaluated in all patients at baseline and after the 4-week intervention.

Limb volume

Upper limb volumes were calculated from circumference measurements taken at 4-cm intervals from the dorsum of the wrist to the axilla. Volume was calculated from circumference using the established formula [21]. Measurements were taken before and after the 4-week intervention by the same therapist [22]. The diagnostic criterion for lymphedema was a ≥ 3 % volume increase from baseline in the affected upper limb [23].

Muscular strength

Shoulder flexor and abductor and elbow flexor muscle strength were evaluated by maximal voluntary isometric contraction using a hand-held dynamometer (Power Track II Commander; JTech Medical, Salt Lake City, UT). Shoulder flexor and abductor testing were performed with the patient in a seated position [24]. Muscular strength was tested in the middle of the joint ROM [24, 25], and the maximum contraction values were used. Maximum isometric contraction was measured over a 5-s duration. A pretest was performed before muscular strength assessment. Patients were allowed a 2-min rest between contractions to avoid fatigue. All muscular strength measurements were performed three times, and the average values were used for analysis.

Range of motion

AROM measurements were performed according to the methods described by Norkin and White [26]. AROM was measured in degrees using a digital inclinometer (The

Saunders Group Inc., Chaska, MN). Upper extremity AROM was evaluated in the following positions: supine flexion and supine abduction. AROM measurements of the affected extremity were performed three times, and the average values were used for analysis. Intra-tester reliability for measurement of AROM was established in a pilot study (intraclass correlation coefficient 3,1=0.84–1.0).

Quality of life

Quality of life was evaluated using the European Organization for Research and Treatment of Cancer Core Quality of Life Questionnaire (EORTC QLQ-C30; version 3) and Breast Cancer-Specific Quality of Life Questionnaire (EORTC QLQ-BR23). The EORTC QLQ-C30 is a 30-item questionnaire consisting of five functional scales (physical, functional, cognitive, emotional, and social performance), three symptoms scales (fatigue, pain, and nausea and vomiting), and scales of quality of life and overall health status. The BR23 is a 23-item questionnaire consisting of functional and symptomatic scales. These questionnaires have been validated and cross-culturally tested in various cancer populations [27].

Arm disability

The disabilities of the arm, shoulder, and hand (DASH) outcome measure is a 30-item questionnaire designed to assess musculoskeletal disorders of the upper limbs. At least 27 questions must be completed for scoring. DASH scores range from 0 to 100, with higher scores indicating a greater level of disability [28]. The DASH has been reported to be a valid and reliable tool for assessing a variety of arm function [29].

Pain

The NRS assesses pain severity using a single 11-point (0 to 10) scale. A score of 0 indicates no pain, whereas a score of 10 indicates severe pain. NRS-based assessments have been shown to provide valid and reliable assessments in patients with cancer [30, 31]. For each assessment, patients were asked to rate the average pain in their arm during abduction of the shoulder, experienced over the past 7 days.

Cording

The presence of visible and palpable cords in the axilla and arm were assessed by a rehabilitation doctor.

Statistical analysis

Statistical analyses were performed using SPSS version 18.0 software (SPSS Inc., Chicago, IL). Continuous and categorical variables are expressed as means±standard deviations and

percentages, respectively. Baseline descriptive statistics were compared using independent *t* tests for continuous data and Fisher's exact test for categorical data. The paired *t* test was used to compare changes in the variables from baseline to week 4 in the two groups. Independent *t* and Fisher's exact tests were applied to compare preintervention to postintervention changes in continuous and categorical variables between the two groups, respectively. A *P* value <0.05 was considered statistically significant.

Results

Comparison of demographic and clinical characteristics between the PTMLD and PT groups

Occupational status, average exercise frequency, educational status, marital status, economic status, chemotherapy, hormone therapy, disease stage, and surgery method were not significantly different between the PTMLD and PT groups. The average patient age was 46.6 years in the PT group and 50.7 years in the PTMLD group. Neither of these differences were statistically significant (Table 1).

Baseline comparisons between the PTMLD and PT groups

In the baseline evaluation, quality of life including functional status (physical, role, emotional, social, and cognitive functioning) and symptoms (fatigue and pain) was not significantly different between the PTMLD and PT groups. Furthermore, arm and breast symptoms (EORTC QLQ-BR23), muscular strength, ROM, DASH score, arm volume, and NRS score were also not significantly different between the groups (Table 2).

Changes in quality of life, muscular strength, AROM, DASH score, NRS score, and arm volume from baseline to the end of the 4-week treatment

Changes in quality of life, muscular strength, AROM, DASH, NRS, and arm volume from baseline to the end of the 4-week treatment are shown in Table 3. EORTC QLQ-C30 and BR23 results showed a significant improvement in physical, role, emotional, and social functioning; fatigue; pain; and arm and breast symptoms in both the PTMLD and PT groups (*P*<0.05). Shoulder flexor strength, shoulder flexion and abduction ROM, DASH score, and pain NRS score were also significantly improved in both groups (*P*<0.05). Arm volume significantly increased over time in the PT group (*P*<0.05).

Table 1 Baseline demographic and clinical characteristics of the patients

Characteristic	PTMLD group (<i>n</i> =21) No. (%) (unless otherwise stated)	PT group (<i>n</i> =20) No. (%) (unless otherwise stated)	<i>P</i> value
Age (years), mean±SD	46.6±6.8	50.7±9.6	0.098
Occupational status			
Non-employed	4 (19)	2 (10)	0.368
Full-time employed	17 (81)	18 (90)	
BMI			
≥25	6 (28.6)	9 (45)	0.500
<25	15 (71.4)	11 (55)	
Exercise frequency			
≥3 times a week for 30 min	11 (52.4)	6 (30)	0.686
<3 times a week for 30 min	10 (47.6)	14 (70)	
Education level			
High school education	3 (14.3)	8 (40)	0.653
University education	18 (85.7)	12 (60)	
Marital status			
Married	17 (81)	18 (90)	0.632
Single	4 (19)	2 (10)	
Economic status/income			
High	4 (19)	4 (20)	0.222
Medium	15 (71.4)	12 (60)	
Low	2 (9.5)	4 (20)	
Chemotherapy			
Yes	9 (42.9)	11 (55)	0.095
No	12 (57.1)	9 (45)	
Radiotherapy			
Yes	21 (100)	19 (95)	0.200
No	0 (0)	1 (5)	
Hormone therapy			
Yes	14 (66.7)	12 (60)	0.187
No	7 (33.3)	8 (40)	
Cancer stage			
I	5 (23.8)	12 (60)	0.019
III	16 (76.2)	8 (40)	
Lesion			
Rt. side	11 (52.4)	11 (55)	0.867
Lt. side	10 (47.6)	9 (45)	
Type of surgery			
Mastectomy	12 (57.1)	16 (80)	0.289
Lumpectomy	7 (33.3)	3 (15)	
Breast reconstruction	2 (9.6)	1 (5)	

BMI body mass index, *PT* physical therapy, *PTMLD* physical therapy combined with manual lymphatic drainage, *SD* standard deviation

Comparison of the incidence of lymphedema and visible cording between the PTMLD and PT groups after the 4-week treatment

Lymphedema occurred in six patients in the PT group and in none of the patients in the PTMLD group ($P<0.05$; Table 4). Visible cording was not significantly different between the groups and was present in 28.5 % of patients in the PTMLD group and 35 % patients in the PT group (Table 4).

Comparison of preintervention to postintervention changes in quality of life, muscular strength, DASH score, NRS score, and arm volume between the PTMLD and PT groups

In the postintervention evaluation, the PTMLD group showed a more significant decrease in NRS score compared with the PT group ($P<0.05$; Table 5). Pain based on the EORTC QLQ-C30 was also significantly decreased in the PTMLD group

Table 2 Baseline comparisons between the PTMLD and PT groups

		PTMLD group (n=21) Mean±SD	PT group (n=20) Mean±SD	P value	
EORTC QLQ-C30	Global health status/QOL	50.8±16.4	48.3±21.6	0.682	
	Functional scales	Physical functioning	69.8±13.9	63.3±17.9	0.200
		Role functioning	68.2±18.2	55.8±30.7	0.128
		Emotional functioning	66.2±11.9	66.6±19.3	0.936
		Cognitive functioning	73.0±14.4	70.2±24.7	0.668
		Social functioning	64.2±27.0	63.3±24.6	0.906
	Symptom scales	Fatigue	42.3±20.9	43.8±15.9	0.791
Pain		43.6±24.4	46.6±22.7	0.685	
EORTC QLQ-BR23	Symptom scales	Breast symptoms	31.3±21.2	26.2±16.5	0.397
	Arm symptoms	43.3±19.5	42.7±22.9	0.929	
Muscular strength (N)	Shoulder flexor	3.8±1.1	3.4±0.8	0.159	
	Shoulder abductor	3.9±1.1	3.5±0.8	0.208	
	Elbow flexor	5.0±1.0	4.7±0.9	0.423	
AROM (degrees)	Shoulder flexion	156.2±9.3	157.3±8.2	0.702	
	Shoulder abduction	150.2±11.1	150.3±7.5	0.997	
DASH score		24.4±16.8	28.3±22.7	0.527	
Arm volume affected region		1895.8±327.2	1847.9±236.1	0.596	
Non-affected region		1895.6±326.5	1846.9±237.6	0.589	
NRS score		6.2±1.4	6.2±1.4	0.841	

AROM active range of motion, EORTC QLQ-C30 European Organization for Research and Treatment of Cancer Core Quality of Life Questionnaire, EORTC QLQ-BR23 European Organization for Research and Treatment of Cancer Breast Cancer-Specific Quality of Life Questionnaire, DASH disabilities of the arm, shoulder, and hand, NRS numeric rating scale, PT physical therapy, PTMLD physical therapy combined with manual lymphatic drainage, QOL quality of life, SD standard deviation

compared with the PT group ($P<0.05$; Table 5). In addition, a significantly greater decrease in arm volume was observed in the PTMLD group ($P<0.05$; Table 5).

Discussion

In our study, pain and arm volume improved in the PTMLD group compared to the PT group. Our hypothesis of this study was that effects of PTMLD group in aspect of shoulder functions would further decrease pain and lymphedema symptoms in comparison to only PT in breast cancer patients with AWS. To the best of our knowledge, this is the first randomized study to evaluate MLD in breast cancer patients with AWS. We chose to evaluate MLD which was the well-known method for lymphedema treatment based on the study by O'Toole et al. [32], which demonstrated that cording of symptom in AWS was an independent risk factor for arm volume increase in breast cancer patients. Furthermore, AWS is a condition reflecting lymph stasis. MLD has been shown to improve blood circulation [18], enhance the movement of lymphatic and other tissue fluids [18], and promote tissue softening and fluid clearance to different and unblocked lymphatic territories [19]. In addition to MLD, active and passive stretching, soft

tissue and cord stretching techniques, and shoulder girdle mobilization were also performed to improve shoulder function. In our results, the cording was not different between the groups. However, MLD affected arm volume in AWS. Our findings support the study by Kepics [33] who recommended MLD, with a 15–20-mmHg compression sleeve if needed, for breast cancer patients with AWS who develop lymphedema.

The pain reduction in the PTMLD group and the difference between the two groups are clinically meaningful. In the present study, patients in the PTMLD group reported significantly less pain than patients in the PT group. AWS patients complained of pain described as a pulling feeling. Previous studies have reported that MLD reduces muscular pain, lymphedema, chronic complex regional pain syndrome, and local levels of inflammatory mediators in subjects with athletic injuries [17]. MLD was also found to relieve pain [33].

Arm lymphedema is the most common complication of breast cancer therapy [18]. Some studies have reported early stimulation of lymphatic drainage is important because inflammation secondary to surgical injuries can render lymphatic drainage difficult [20, 34, 35].

Torres Lacomba et al. [20] suggested that MLD can reduce the risk of lymphedema 2 years after surgery. In our study, lymphedema occurred in the PT group, but not in the

Table 3 Changes in QOL, muscular strength, AROM, DASH score, NRS score, and arm volume from baseline to the end of the 4-week treatment

			Baseline Mean±SD	After 4-week treatment Mean±SD	Mean difference (95 % CI)	<i>P</i> value	
EORTC QLQ-C30	Global health status/QOL	PTMLD	50.8±16.4	66.2±16.6	−15.48 (−22.40 to −8.56)	0.000	
		PT	48.3±21.6	61.2±21.0	12.95 (5.09 to 20.80)	0.003	
	Functional scales	Physical functioning	PTMLD	69.8±13.9	79.0±10.2	−9.21 (−14.34 to −4.09)	0.001
			PT	63.3±17.9	73.6±15.4	10.33 (3.97 to 16.69)	0.003
		Role functioning	PTMLD	68.2±18.2	83.3±17.5	−15.09 (−23.02 to −7.16)	0.001
			PT	55.8±30.7	62.5±22.2	6.66 (0.78 to 12.54)	0.028
		Emotional functioning	PTMLD	66.2±11.9	81.3±13.2	−15.09 (−21.26 to −8.91)	0.000
			PT	66.6±19.3	77.1±16.2	10.41 (3.98 to 16.85)	0.003
	Cognitive functioning	PTMLD	73.0±14.4	84.9±12.8	−11.92 (−18.78 to −5.06)	0.002	
		PT	70.2±24.7	84.1±14.8	13.92 (3.61 to 24.23)	0.011	
	Social functioning	PTMLD	64.2±27.1	85.7±16.1	−21.46 (−34.65 to −8.27)	0.003	
		PT	63.3±24.6	76.6±14.7	13.37 (2.13 to 24.60)	0.022	
	Symptom scales	Fatigue	PTMLD	42.3±20.9	23.8±20.9	18.50 (10.61 to 26.38)	0.000
			PT	43.8±15.9	30.5±17.2	−13.32 (−25.19 to −1.44)	0.030
		Pain	PTMLD	43.6±24.4	13.5±14.5	30.15 (23.10 to 37.19)	0.000
			PT	46.6±22.7	30.0±16.8	−16.66 (−24.25 to −9.06)	0.000
EORTC QLQ-BR23	Symptom scales	Breast symptoms	PTMLD	31.3±21.2	10.7±11.5	20.63 (12.44 to 28.82)	0.000
			PT	26.3±16.9	8.8±14.3	−17.53 (−25.66 to −9.40)	0.000
	Arm symptoms	PTMLD	43.3±19.5	23.3±15.7	20.07 (12.47 to 27.66)	0.000	
		PT	42.7±22.9	19.4±15.6	−23.31 (−34.47 to −12.14)	0.000	
Muscular strength (N)	Shoulder flexor	PTMLD	3.8±1.1	4.0±1.0	−0.14 (−0.27 to −0.01)	0.028	
		PT	3.4±0.8	3.5±0.8	0.13 (0.02 to 0.23)	0.023	
	Shoulder abductor	PTMLD	3.9±1.1	4.0±0.9	−0.1 (−0.25 to 0.04)	0.153	
		PT	3.5±0.8	3.7±0.8	0.18 (0.04 to 0.31)	0.013	
	Elbow flexor	PTMLD	4.9±1.0	5.1±0.8	−0.13 (−0.38 to 0.11)	0.260	
		PT	4.7±0.9	4.7±0.8	−0.02 (−0.15 to 0.11)	0.765	
AROM (degrees)	Shoulder flexion	PTMLD	156.2±9.3	180.0±0.0	−23.80 (−28.06 to −19.55)	0.000	
		PT	157.3±8.2	180.0±0.0	22.75 (18.91 to 26.58)	0.000	
	Shoulder abduction	PTMLD	150.2±11.1	180.0±0.0	−29.76 (−34.82 to −24.69)	0.000	
		PT	150.3±7.5	180.0±0.0	29.75 (26.23 to 33.26)	0.000	

Table 3 (continued)

		Baseline Mean±SD	After 4-week treatment Mean±SD	Mean difference (95 % CI)	<i>P</i> value
DASH score	PTMLD	24.4±16.8	15.7±10.6	8.66 (4.48 to 12.83)	0.000
	PT	28.3±22.7	14.2±9.8	-14.11 (-21.28 to -6.93)	0.001
Arm volume	PTMLD	1895.8±327.2	1894.7±326.7	1.09 (-0.38 to 2.57)	0.139
	PT	1847.9±236.1	1869.7±242.8	21.75 (1.58 to 41.91)	0.036
NRS score	PTMLD	6.2±1.4	1.5±1.0	4.76 (4.01 to 5.42)	0.000
	PT	6.2±1.4	2.6±1.3	-3.6 (-4.11 to -3.08)	0.000

AROM active range of motion, *EORTC QLQ-C30* European Organization for Research and Treatment of Cancer Core Quality of Life Questionnaire, *EORTC QLQ-BR23* European Organization for Research and Treatment of Cancer Breast Cancer-Specific Quality of Life Questionnaire, *DASH* disabilities of the arm, shoulder, and hand, *NRS* numeric rating scale, *PT* physical therapy, *PTMLD* physical therapy combined with manual lymphatic drainage, *QOL* quality of life, *SD* standard deviation, *CI* confidence interval

PTMLD group. Six of the 20 patients in the PT group developed lymphedema after 4 weeks. The affected arm had 3 % difference in volume and patients were diagnosed secondary lymphedema by using lymphoscintigraphic findings, which showed decreased function of the affected axilla lymph nodes. Three of them had increased uptake through dermal backflow. Patients in the PT group also showed a significant increase in arm volume from baseline to week 4 of treatment. Reported lymphedema incidence rates range from 13 to 65 % among breast cancer survivors, depending on the diagnostic criteria used [36, 37]. Our 33.3 % lymphedema incidence rates consistent with previous reports. Because the PTMLD group did not have increase of arm volume, we think our intervention did not cause lymphedema. Also, there were no previous studies that demonstrate physical therapy can cause lymphedema in AWS.

AWS usually occurs in the early postoperative period after breast cancer surgery and is a painful condition that limits daily life functioning. Shoulder abduction is particularly painful in patients with AWS. The present study included patients at least 4 weeks after surgery. And most of them were included at early periods after surgery. The tendency of patients to protect the surgical area and AWS-induced pain can progressively limit shoulder mobility. Previous studies have reported that AWS resolves spontaneously within 3 months [2, 10].

However, during this period of time, changes in shoulder girdle movement and muscle imbalances may occur. We found that AROM (flexion, abduction) of the shoulder was significantly improved in both the PTMLD and PT groups following treatment. In addition, the physical therapy program increased shoulder flexor muscle strength and DASH score. Josenhans [38] demonstrated that symptom improvement including decreased pain, restoration of full shoulder ROM, and increased shoulder function could be attained within six physical therapy sessions. In our study, physical therapy restored shoulder ROM and improved muscular strength in all patients. Most studies of physical therapy in breast cancer patients with AWS have been case reports. Nonetheless, these studies suggest that physical therapy is a quick and effective method to resolve shoulder disorders [11, 12]. Fourie and Robb [39] reported the case of a woman who developed AWS 22 days following axillary dissection. Physiotherapy with rotatory movements and stretching of the restricted tissue resulted in the achievement of pre-morbid ROM within 3 weeks. Furthermore, pain and visible cords had completely resolved after 16 weeks of therapy. As this case report, physical therapy can shorten the natural course of AWS by up to 3 months.

Breast cancer patients commonly experience symptoms of fatigue, anxiety, and depression. Breast cancer and its

Table 4 Comparison of the incidence of lymphedema and visible cording between the PTMLD and PT groups after 4 weeks of treatment

	PTMLD group (<i>n</i> =21) Mean±SD	PT group (<i>n</i> =20) Mean±SD	Difference	<i>P</i> value
Visible cords (yes/no)	6/15	7/13	1	0.658
Lymphedema (yes/no)	0/21	6/14	6	0.009

PT physical therapy, *PTMLD* physical therapy combined with manual lymphatic drainage, *SD* standard deviation

Table 5 Comparison of preintervention to postintervention changes in QOL, muscular strength, AROM, DASH score, NRS score, and arm volume between the PTMLD and PT groups

		PTMLD group (n=21) Mean±SD	PT group (n=20) Mean±SD	Mean difference (95 % CI)	P value	
EORTC QLQ-C30	Global health status/QOL	15.5±15.2	13.0±16.8	2.53 (−7.60 to 12.66)	0.615	
	Functional scales	Physical functioning	9.2±11.3	10.3±13.6	−1.11 (−9.03 to 6.79)	0.776
		Role functioning	15.1±17.4	6.7±12.6	8.43 (−1.15 to 18.01)	0.085
		Emotional functioning	15.1±13.6	10.4±13.8	4.67 (−3.96 to 13.30)	0.281
	Cognitive functioning	11.9±15.1	13.9±22.0	−2.00 (−14.04 to 10.04)	0.735	
	Social functioning	21.5±29.0	13.3±24.0	8.09 (−8.68 to 24.88)	0.337	
	Symptom scales	Fatigue	−18.5±17.3	−13.3±25.4	−5.18 (−19.03 to 8.67)	0.448
Pain		−30.2±15.5	−16.7±16.2	−13.49 (−23.52 to −3.46)	0.010	
EORTC QLQ-BR23	Symptom scales	Breast symptoms	−20.6±18.0	−17.9±16.5	−2.72 (−13.62 to 8.17)	0.616
		Arm symptoms	−20.1±16.7	−23.3±23.9	3.23 (−9.89 to 16.36)	0.616
Muscular strength (N)	Shoulder flexor	0.1±0.3	0.1±0.2	0.01 (−0.14 to 0.18)	0.831	
	Shoulder abductor	0.1±0.3	0.2±0.3	−0.07 (−0.27 to 0.12)	0.442	
	Elbow flexor	0.1±0.5	0.0±0.3	0.15 (0.11 to 0.43)	0.259	
AROM (degrees)	Shoulder flexion	23.8±9.3	22.8±8.2	1.06 (−4.48 to 6.60)	0.702	
	Shoulder abduction	29.8±11.1	29.8±7.5	0.01 (−5.98 to 6.00)	0.997	
DASH score		−8.7±9.2	−14.1±15.3	5.44 (−2.65 to 13.54)	0.180	
Arm volume		−1.1±3.3	21.8±43.1	−22.84 (−41.90 to −3.78)	0.029	
NRS score		−4.8±1.4	−3.6±1.1	−1.16 (−1.97 to −0.34)	0.006	

AROM active range of motion, EORTC QLQ-C30 European Organization for Research and Treatment of Cancer Core Quality of Life Questionnaire, EORTC QLQ-BR23 European Organization for Research and Treatment of Cancer Breast Cancer-Specific Quality of Life Questionnaire, DASH disabilities of the arm, shoulder, and hand, NRS numeric rating scale, PT physical therapy, PTMLD physical therapy combined with manual lymphatic drainage, QOL quality of life, SD standard deviation, CI confidence interval

treatment pose many challenges to physical, emotional, mental, and social well-being and negatively impact quality of life. In particular, limited shoulder ROM negatively influences functional capacity and quality of life [40–42]. Breast cancer patients with musculoskeletal pain demonstrate significantly lower health-related quality of life including physical and mental functioning [41]. Therefore, QoL is an important patient-centered outcome addressed by PT in breast cancer patients. Indeed, quality of life was improved in both the PTMLD and PT groups in our study. The physical therapy program improved physical, role, emotional, and social functioning and fatigue in the study population. Furthermore, exercise had a positive effect on breast and arm symptoms. Our

physical therapy program focused on shoulder stretching exercises. We believe that restoration of shoulder function and decreased pain contributed to the positive outcomes of the physical therapy program on quality of life. Previous studies have shown that yoga programs also improve stress, fatigue, emotional functioning, pain, vitality, and quality of life in cancer patients [42, 43].

Our study had several limitations. First, we could not discern the natural course of lymphedema development and AWS. The lack of a true, no-treatment, control group prevented us from observing the natural course of AWS. A no-treatment control group was not included because we considered it unethical to withhold treatment from patients with

AWS. Second, we did not consider of change of arm volume by assessing muscle hypertrophy through imaging but relied solely on circumferential assessment. However, this is the commonly used clinical assessment so has clinical relevance. Third, the PT MLD group received additional PT time and attention during the first week, which could be a potential confounder and may have had an impact on reports of pain. Finally, patient compliance in taking prescribed NSAIDs was not monitored. In conclusion, our findings provide evidence that a physical therapy program combined with MLD is an effective strategy to improve pain, shoulder function, and QOL in breast cancer patients with AWS. Physical therapy can also contribute to the prevention of lymphedema and shoulder joint contracture due to pain.

Compliance with Ethical Standards

Conflict of interest None of the authors has any financial support or relationship with regard to the submitted manuscript that might be construed as a conflict of interest. All authors have full control of all primary data and agree to allow the journal to review the data if requested.

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