

CURRENT ROLE OF PNEUMATIC COMPRESSION THERAPY IN LYMPHEDEMA CARE: A SCOPING REVIEW OF PERSISTENT DEBATES AND NEW APPLICATIONS

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ABSTRACT

Access to trained lymphedema care providers remains limited making patient-driven management solutions essential. One such option, sequential intermittent pneumatic compression (IPC), has gained traction as a supportive tool for lymphedema management. While newer IPC devices and innovative applications are being introduced to the market, questions regarding the safety and efficacy of this technology persist. This underscores the importance of reviewing current literature to understand IPC's evolving role in lymphedema care and to identify existing knowledge gaps. A scoping review of literature was conducted across various databases using PRISMA-ScR guidelines. The eligibility criteria included articles published in English language from database's inception to June 2023, discussing IPC's safety, and/or efficacy, and/or optimal modes and settings for lymphedema management. Review articles and case reports and original studies with unclear outcome measures were excluded. The review identified 49 eligible studies from an initial pool of 614 articles, consisting of 12 randomized controlled trials, 25 cohort studies, and 12 experimental studies. Most studies (44) focused on limb lymphedema, while five examined non-limb regions. Sample sizes varied widely, ranging from 10 to 718 participants, reflecting differences in studies' power. Minor adverse events

were reported in six studies, including transient skin irritation, paresthesia, and rare cases of genital edema. Efficacy data indicated that IPC, whether used with or without manual lymphatic drainage (MLD), improved limb volume, quality of life, and reduced infection rates, although results varied according to treatment protocols and limb type. The addition of IPC improved compliance of decongestive therapy and increased patient satisfaction. IPC sessions ranged from 45 to 120 minutes per day, conducted 3 to 7 days per week, with pressures set at 60 to 120 mmHg for lower limbs and 25 to 60 mmHg for upper limbs. Higher pressures were associated with more significant limb volume reduction in the lower limbs. A cost analysis indicated that IPC could potentially lead to healthcare savings by reducing infections and hospital admissions. IPC application also showed promising results in head and neck lymphedema, though results for trunk lymphedema were equivocal. Future research should aim to refine IPC protocols in different regions of the body and ascertain its long-term benefits.

Keywords: Intermittent Pneumatic Compression, Lymphedema, Best Practices, Safety, Review

Lymphedema is characterized by chronic accumulation of lymphatic fluid, fat, and fibrosis which presents a complex therapeutic

challenge. Complete decongestive therapy (CDT), encompassing manual lymphatic drainage (MLD), compression, exercise, and skin care, remains the cornerstone of conservative management. Although there have been significant advancements in lymphatic surgery, elevating the goals of modern lymphedema management, access to these specialized surgical interventions, as well as to trained lymphedema therapists remains limited. This dual constraint necessitates patient-driven self-management. Intermittent pneumatic compression (IPC) pump has emerged as a viable at-home supplement to traditional lymphedema treatments, empowering patients with greater control over their condition (1). The market offers a variety of IPC devices, each differing in sleeve and pump design, configuration, programmability, chamber count, inflation/deflation cycles, maximum pressure, and pressure gradient. The advent of new compression modes, such as those mimicking MLD, further complicates the landscape. Recently, IPC usage has expanded to include other body areas, such as the head and neck, highlighting the need for comprehensive review. Recent innovations have further expanded IPC capabilities, including applications for non-limb areas such as the head and neck and introduction of new compression modes. While previous reviews have explored IPC effectiveness, they haven't addressed the recent advancements. Also, experimental studies using imaging to optimize compression settings have not been reviewed (2-5). This literature review aims to fill these gaps by synthesizing current evidence on IPC usage in lymphedema management and identifying areas requiring further research.

METHODS

Search Strategy and Information Sources

We conducted a scoping literature review following PRISMA-ScR guidelines utilizing Ovid MEDLINE, PubMed, EMBASE, and the Cochrane Library databases covering the period from their inception to June 2023 (6). The search was limited to articles published in

English language. Keywords and MeSH terms included "intermittent pneumatic compression", OR "compression therapy", OR "compression pumps", OR "sequential compression" AND "lymphedema" OR "lymphatic flow". Cross references of the selected articles were screened for any relevant articles and included in the search results.

Eligibility Criteria

Inclusion criteria included studies evaluating safety and/or efficacy and/or optimal machine settings of IPC in lymphedema patients. Eligible studies encompassed both pediatric and adult patients with lymphedema affecting the extremities, trunk, or head/neck. Primary outcomes of interest were imaging findings, changes in limb volume or circumference, patient-reported outcomes, quality of life assessments, and adverse events. Case reports, literature reviews, and studies with unclear outcome measures were excluded. Also, studies focusing on non-lymphedema patients, or animal studies, were eliminated.

Study Selection and Data Extraction

Four independent reviewers conducted a screening process of titles, abstracts, and full texts using a web-based tool, Covidence (Melbourne, Australia). Studies that met the eligibility criteria were identified individually and any discrepancies were resolved through consensus. Data extraction was carried out using a standardized form generated in Covidence, capturing information on study design, study aim, cohort characteristics, sample size, lymphedema etiology, affected regions, outcome measures, treatment protocols, IPC device specifications, duration of study, and key findings.

Synthesis of Results

We categorized the outcomes into following domains – safety, efficacy, optimal device settings, and cost effectiveness. The efficacy results were subcategorized based on evaluation methodology, the treated region,

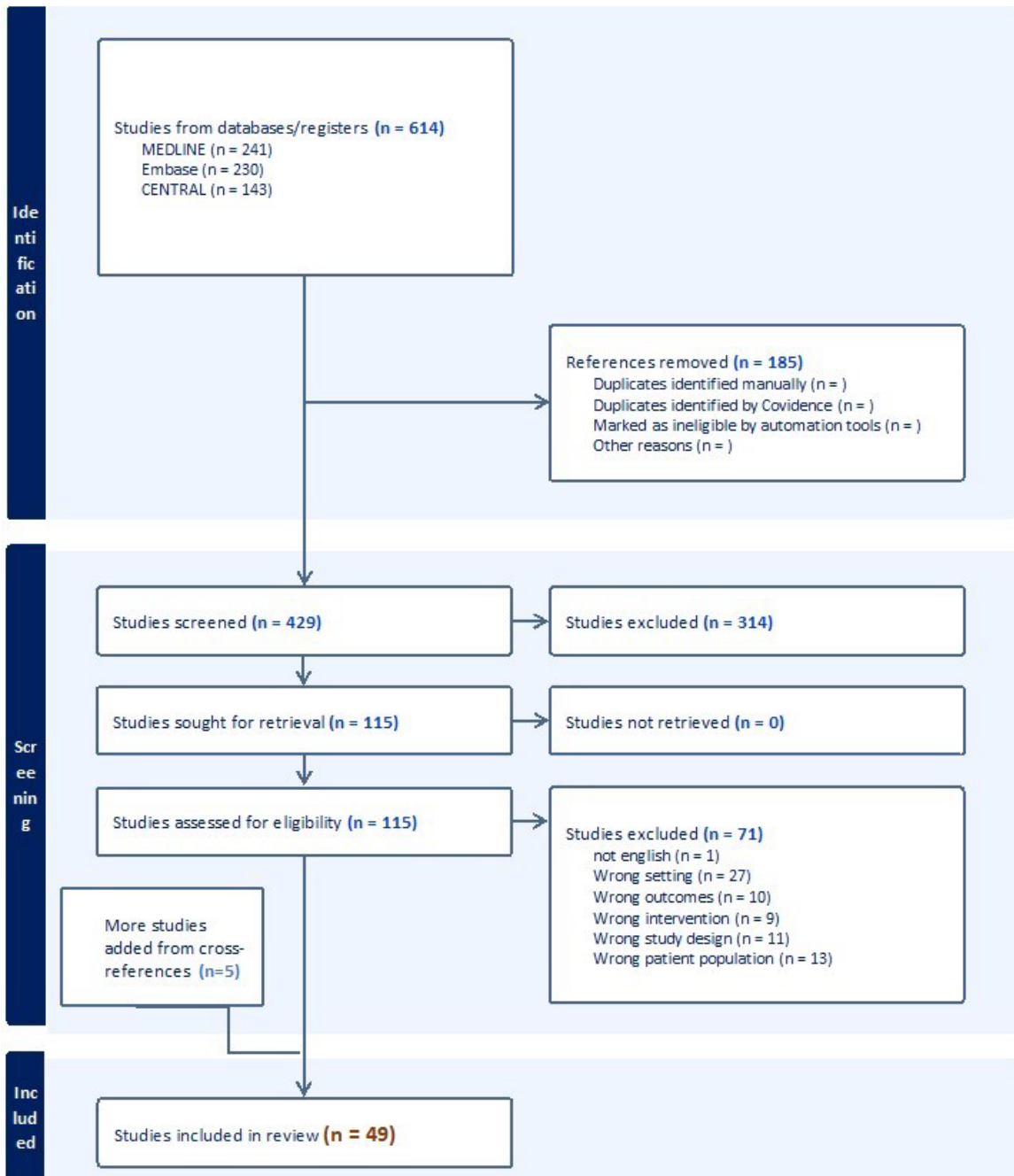


Fig. 1. PRISMA diagram detailing study flow from initial manuscripts to final 49 utilized for the scoping review.

and any special situations. Evidence for sustainability of effects, variations based on the affected limb, and efficacy of new compression modes were also reviewed.

RESULTS

Study Characteristics

The search yielded 614 articles, of which 49 studies satisfied eligibility criteria (*Fig. 1*). *Table 1* (larger version in supplementary materials) presents the characteristics of the included studies. These encompassed 12 randomized controlled trials, 18 observational studies, 7 case series and 12 experimental studies. Forty-four articles focused on lymphedema in the limbs, while five studies explored non-limb regions, including the head, neck, and trunk. The sample sizes in the studies varied significantly, ranging from 10 to 718 patients, reflecting the diversity in the power of the studies. The findings of the studies are presented below.

Safety

IPC is contraindicated in patients with underlying peripheral arterial disease, active deep venous thrombosis, active infections, and/or evidence of cancer recurrence (2). There is lack of consensus on the safety of IPC in patients with cardiac failure where the severity of cardiac dysfunction should be considered to weigh the risks and benefits. Vigilant monitoring is essential to circumvent potential cardiac decompensation in patients with severe disease as a result of augmented venous return (7).

We screened all selected studies for report on adverse events associated with IPC use in patients with lymphedema. Only six studies provided information on adverse events in their study population (1,8-12). In these studies, minimal effects like transient skin irritation, paresthesia, mild pain (25% patients), muscle cramps, and limb erythema were noted. Patient discomfort attributable to IPC was ameliorated by modulation of the applied pressures (9). Rare events included headache, nausea, and dizziness (10). One

particular study highlighted a significant disparity in genital edema incidence, registering a 43% occurrence with IPC intervention as opposed to a 3% occurrence without IPC (12). The duration for which the genital edema lasted was not delineated in the study. No other study reported increase in genitalia swelling although mild groin and knee swelling was reported in another study which was transient and could be mitigated by additional groin compression (13). In aggregate, the studies under review did not report any major complications ensuing from IPC treatment.

Efficacy

Total 37 studies examined the efficacy of IPC using various strategies. The efficacy of IPC in limb lymphedema was assessed in comparison to manual lymphatic drainage (MLD), as an adjunct to MLD, or as a standalone treatment (no MLD). All studies examining the absolute benefit of IPC as a standalone therapy allowed patients to continue using compression garments by default, hence was not in effect a solitary management tool. The most common outcome measure used for determining efficacy was limb circumference or volume change. Patient-reported outcomes, such as change in symptoms, range of motion, quality of life, reduction in cellulitis episodes, and hospital admission rates was utilized by several studies (*Table 1*). Six experimental studies used imaging techniques including ICG lymphography or lymphoscintigraphy to objectively study the immediate effects of pneumatic compression on lymphatic flow after a single treatment session (14-18). The only prospective clinical study that utilized imaging was conducted in head and neck lymphedema cohort, using ICG lymphography before and after two-weeks of IPC therapy (19).

IPC

Eight studies evaluated IPC as a stand-alone therapy (without MLD but with compression garments) for primarily lower limb (7 studies, primary and acquired disease) or

TABLE 1
Detailed Manuscript Information Grouped into 4 Groups used for Analysis
(Limb Lymphedema, Special Situations, Non-Limb Lymphedema,
and Relevant Compression Pressure Studies)

Author, Year	Title	Summary of Outcomes	Salient point	Sample size	Enrichy	Region	Total Duration of	Duration per Day	Max Follow up period	Frequency	Compression Cycles	IPC Pressure	Device name	Study Design	Outcome measures	Adverse events
Enayati 2022	Severe lymphedema in gynecological cancer: A retrospective study of the effect of lymphatic decongestive treatment on the quality of life of patients with lymphedema	Following IPC, participants showed a significant improvement in mean AFCD score. The EFMQOL scores showed a significant improvement in the AFCD score of patients with lymphedema.	IPC improves all relevant functional and symptom scores, except for a significant limb girth reduction at 12 weeks.	12	secondary	LE	8w	1h	N/A	daily	N/A	100 mmHg	MA	Prospective cohort study	Quality of life scores and history of lymphedema	
Maldonado 2021	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	The first one-hour IPC showed decrease in circumference of lower parts of the calf and thigh with increase below knee and in the AFCD. The AFCD treatment was associated with improvement of a wide range of IPC with significant reductions in LY and showed that IPC can be used in chronic cases of the lymphedema.	74	secondary	LE	52w	45m	52 weeks	5 days per week	N/A	N/A	FlexTouch	Prospective cohort study	The primary endpoint: OCL change at 12 weeks. Secondary endpoints: EFMQOL score, number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Dress 2019	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	117	secondary	UE, LE	52w	1h	52 weeks	twice daily	18 s per cycle	40 to 60 mmHg	Medi Pump, Corder	Prospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Blumberg 2016	Improvement in the quality of life in patients with lymphedema after treatment with IPC	The use of IPC was associated with a significant decrease in the use of hospitalizations, patient satisfaction with treatment, and patient compliance.	IPC may have limited clinical value.	100	secondary	LE	51w	45m	51 weeks	5 to 7 days per week	N/A	N/A	MA	Prospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Byrnes 2014	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	1065	secondary	N/A	N/A	N/A	N/A	N/A	N/A	N/A	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Zubala 2014	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	18	secondary	LE	2 to 3 years	45 min	3 months	3 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Mohr 2013	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	196	secondary	LE	N/A	N/A	4 to 9 weeks	daily	N/A	80 to 110 mmHg	N/A	Prospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
McLaughlin 2010	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	45	secondary	LE	2 days	8h	2 months	5 days per week	Sequential	80 to 120 mmHg	N/A	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Borns 1998	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	128	secondary	LE	6w	2h	3 months	3 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Guzo 1996	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	67	secondary	UE	5w	2h	3 months	3 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Stahl 1994	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	46	secondary	UE	6w	45m	2 months	5 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Ortiz 2012	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	30	secondary	UE	4w	45m	2 months	5 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Fonseca 2021	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	132	secondary	UE, LE	6w	30m	2 months	5 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Johnson 2021	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	28	secondary	UE	2w	2 hrs	3 months	5 days per week	Sequential	40-60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Swan 2022	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	69	secondary	LE	78w	N/A	2 months	5 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Sozonsky 2022	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	27 (13 CDT, 14 MLD)	secondary	UE	2w	60 min MLD vs 30 min MLD + 30 min IPC	2 months	5 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Wakabayashi 2020	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	35 CDT, 10 IPC	secondary	UE	4w	60 min MLD vs 60 min MLD + 30 min IPC	2 months	5 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Uzarski 2015	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	31	secondary	UE	3w	45 min	2 months	5 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Uzarski 2013	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	25	secondary	UE	3w	N/A	2 months	5 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Hajkhalil 2010	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	112	secondary	UE	12w	1h	12w	5 days per week	Sequential	60 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Scuba 2002	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	23	secondary	UE	26-52 weeks	30 to 60 minutes	12 months	5 days per week	Sequential	40 to 50 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Tsushiki 2013	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	81	Phakolymp	LE	4w	45m	4 weeks	twice daily	N/A	40 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Korley 2023	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	21	primary, secondary	LE	5 days or 12 days	Group A: 1 hour per day so 2 hours total. Group B: 2 hours (out twice per day so 4 hours total).	12 months	twice daily for 4 weeks	N/A	40 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Drain 2002	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	40	secondary	LE	5w	N/A	12 months	twice daily for 4 weeks	N/A	40 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Kim 2022	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	30	secondary	LE	N/A	1h	12 months	twice daily for 4 weeks	N/A	40 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	
Kenworthy 2017	Effectiveness of the use of IPC in the treatment of lymphedema in breast cancer patients	IPC treatment significantly improved the EFMQOL scores, showed a significant improvement in the AFCD score of patients with lymphedema.	IPC may have limited clinical value.	1731	secondary, primary	unspecified	12 months	N/A	12 months	twice daily for 4 weeks	N/A	40 mmHg	MA	Retrospective cohort study	Primary endpoint: EFMQOL score. Secondary endpoints: number of ulcers, venous number of ulcers, venous rate of hospitalizations, patient satisfaction with treatment, and patient compliance.	

Device name and device	Comparing 2 devices	Feb 2012	A randomized controlled trial comparing two types of pneumatic compression for the treatment of lymphedema in the home	The AFD-DC-based group showed a significant 29% reduction in edema compared to a 16% increase in the IPC control group. The AFD-DC-based group had a 10% increase in the AFD-DC-based group and a 13.1% increase in the AFD-DC-based group. The AFD-DC-based group provides better outcome than with an IPC, and both groups demonstrated very good device compliance.	Advanced IPC as an adjunct treatment may be better than standard IPC device for testing BCEL. Seven patients had adverse events, more in the 5th EP	36	secondary	UE	12w	1h	N/A	7 days per week	18 per chamber (daily sequential) or 9 IPCs (intermittent) each with 13.14±4.8 mmHg and 13.14±4.8 mmHg all inflated vs all inflated vs 1 to 3 ses per chamber (onlyone inflated at a time)	30 mmHg (standard) or 4.2 mmHg (intermittent) and 13.14±4.8 mmHg (Advanced)	N/A	RCT	Limb volume (with circumference and tissue water measurement) (with Mousa and Work 0 and 1, 2)	Seven patients had adverse events, more in the standard group.
Think compression device	Think compression device	Fisher 2012	A within limb, within limb, pneumatic advanced pneumatic manual, chest, and arm treatment to arms treatment only in self-care of lymphedema therapy	When comparing equivalent pneumatic compression therapy to pneumatic compression (control), there was no statistically significant change in IPC is an effective method of volume reduction in women with postmastectomy lymphedema regardless of cycle time? Secondary endpoints: chest, arm, and limb volume; patient compliance; and quality of life.	1h in UE + limb, 36 mins in UE only EP	42	secondary <td>UE <td>30 days</td> <td>N/A</td> <td>7 days per week</td> <td>N/A</td> <td>N/A</td> <td>9.0 ± 4.2-13.7 ± 4.9 mmHg</td> <td>N/A</td> <td>RCT</td> <td>USDS 4-way arm circumference, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume</td> </td>	UE <td>30 days</td> <td>N/A</td> <td>7 days per week</td> <td>N/A</td> <td>N/A</td> <td>9.0 ± 4.2-13.7 ± 4.9 mmHg</td> <td>N/A</td> <td>RCT</td> <td>USDS 4-way arm circumference, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume</td>	30 days	N/A	7 days per week	N/A	N/A	9.0 ± 4.2-13.7 ± 4.9 mmHg	N/A	RCT	USDS 4-way arm circumference, tank circumference, FASCO score, bioimpedance	Limb volume
No of chambers and cycle time	Think compression device	Fisher 2009	Influence of compression cycle time and number of chambers on upper extremity lymphedema	IPC is an effective method of volume reduction in women with postmastectomy lymphedema regardless of cycle time? Secondary endpoints: chest, arm, and limb volume; patient compliance; and quality of life.	1h	37	secondary <td>UE <td>5w</td> <td>Single session for each device</td> <td>Single session for each device</td> <td>Single session for each device</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>RCT</td> <td>Limb volume</td> <td>Limb volume (water displacement)</td> </td>	UE <td>5w</td> <td>Single session for each device</td> <td>Single session for each device</td> <td>Single session for each device</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>RCT</td> <td>Limb volume</td> <td>Limb volume (water displacement)</td>	5w	Single session for each device	Single session for each device	Single session for each device	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	RCT	Limb volume	Limb volume (water displacement)
Comparing 2 devices	Think compression device	Mansueti 2007	Effect of pneumatic compression therapy on lymphedema	These were significant differences between the Phrotouch and Lympho Press devices. During inflation cycles, the pressure exerted by the Lympho Press was higher and was sustained for longer periods of time. However, the Phrotouch deflated more quickly displaying quick rise and fall progression.	2 hrs	35	secondary, primary <td>UE, LE</td> <td>N/A</td> <td>N/A</td> <td>N/A</td> <td>Single session for each device</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	UE, LE	N/A	N/A	N/A	Single session for each device	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
No. of chambers	Think compression device	Bergan 1996	A comparison of the effectiveness of the treatment of lymphedema	One and three chamber garments were less effective than four chambers reducing swelling.	N/A	N/A	secondary, primary <td>UE, LE</td> <td>N/A</td> <td>2 hrs</td> <td>N/A</td> <td>Single session for each device</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>RCT</td> <td>Limb volume (water displacement)</td> <td>Limb volume (water displacement)</td>	UE, LE	N/A	2 hrs	N/A	Single session for each device	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	RCT	Limb volume (water displacement)	Limb volume (water displacement)
Comparing 2 devices	Think compression device	Zahedi 2019	The Effectiveness of Advant Compression Therapy on Lymphedema: A Systematic Review, Meta-Analysis, and Evidence-Based Practice Guidelines	Adjustment of compression parameters to these findings, that accumulation of lymph fluid in the limb is a result of various limb levels is responsible for effective therapy. The recommended differential compression pressure and prolonged hangings at various limb levels	LE	23	secondary, primary <td>LE</td> <td>1 hr</td> <td>N/A</td> <td>Single session for each device</td> <td>Single session for each device</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	LE	1 hr	N/A	Single session for each device	Single session for each device	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
Effectiveness Best settings	Think compression device	Bok 2018	Evaluation of Suffices in Postmastectomy Lymphedema: Effect of compression therapy on lymph movement during and after IPC in all affected leg of the subjects treated with lymph movement in	After a single session of IPC, using a pressure of 35 mmHg resulted in the effective treatment of postmastectomy lymphedema. The mean difference in limb volume during and after IPC in all affected leg of the subjects treated with lymph movement in	N/A	45	secondary <td>UE</td> <td>single session</td> <td>N/A</td> <td>N/A</td> <td>Single session for each device</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	UE	single session	N/A	N/A	Single session for each device	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
Effectiveness Best settings	Think compression device	Ahmed 2017	Effect of compression therapy on lymph movement during and after IPC in all affected leg of the subjects treated with lymph movement in	IPC improves lymphatic function either by modulating fluid through lymph vessels or through interstitial channels.	1h	4	secondary, primary <td>LE</td> <td>single session</td> <td>N/A</td> <td>N/A</td> <td>Single session for each device</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	LE	single session	N/A	N/A	Single session for each device	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
Effectiveness Best settings	Think compression device	Kithama 2017	Real-Time Direct Evidence of the Superiority of Lymphatic Compression in Lymphedema: A Randomized Trial	Different inflation/deflation modes and two different pressures evaluated (45 and 30 mmHg) using quantitative ICG and software-assisted video analysis (EG IntraMotion Pro-united).	single session	27 (8 control, 17 limb)	secondary <td>LE</td> <td>single session</td> <td>single session</td> <td>single session</td> <td>single session</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	LE	single session	single session	single session	single session	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
Effectiveness Best settings	Think compression device	Zahedi 2013	Pressure and timing of compression device for lymphedema	The study aims to measure the effect of different inflation/deflation modes and two different pressures evaluated (45 and 30 mmHg) using quantitative ICG and software-assisted video analysis (EG IntraMotion Pro-united).	single session	13	secondary, primary <td>LE</td> <td>single session</td> <td>single session</td> <td>single session</td> <td>single session</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	LE	single session	single session	single session	single session	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
Effectiveness Best settings	Think compression device	Ossewsky 2011	Tissue fluid pressure and flow during pneumatic compression in lymphedema of lower limb	The study aims to measure the effect of different inflation/deflation modes and two different pressures evaluated (45 and 30 mmHg) using quantitative ICG and software-assisted video analysis (EG IntraMotion Pro-united).	single session	15	N/A <td>LE</td> <td>single session</td> <td>single session</td> <td>single session</td> <td>single session</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	LE	single session	single session	single session	single session	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
Effectiveness Best settings	Think compression device	Ossewsky 2011	Pathways of lymph and tissue fluid flow during compression in lymphedema of lower limb	The study aims to measure the effect of different inflation/deflation modes and two different pressures evaluated (45 and 30 mmHg) using quantitative ICG and software-assisted video analysis (EG IntraMotion Pro-united).	single session	15	secondary <td>LE</td> <td>single session</td> <td>N/A</td> <td>N/A</td> <td>single session</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	LE	single session	N/A	N/A	single session	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
Effectiveness Best settings	Think compression device	Adami 2010	Effect of pneumatic compression therapy on lymphedema	The study aims to measure the effect of different inflation/deflation modes and two different pressures evaluated (45 and 30 mmHg) using quantitative ICG and software-assisted video analysis (EG IntraMotion Pro-united).	single session	3 patients of limb)	secondary <td>UE</td> <td>single session</td> <td>single session</td> <td>single session</td> <td>single session</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	UE	single session	single session	single session	single session	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
Effectiveness Best settings	Think compression device	Manjiv Hassali 2001	Evaluation of sequential pneumatic compression therapy on lymphedema	12 patients with grade II distal lymphedema had >50% reduction in edema.	6 months	28	secondary <td>LE</td> <td>4w</td> <td>N/A</td> <td>single session</td> <td>single session</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	LE	4w	N/A	single session	single session	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)
Effectiveness Best settings	Think compression device	Hassali 2001	In this retrospective study of 10 children with lymphedema of the upper or lower extremities, the lymphedema was	pressure was required decreased over time in the same pt	N/A	16	secondary, primary <td>UE, LE</td> <td>N/A</td> <td>N/A</td> <td>single session</td> <td>single session</td> <td>Sequential of sequential sequential</td> <td>30 mmHg in chambers 50 to 30 mmHg gradient in chamber</td> <td>N/A</td> <td>Experimental study</td> <td>bioimpedance, tank circumference, FASCO score, bioimpedance</td> <td>Limb volume (water displacement)</td>	UE, LE	N/A	N/A	single session	single session	Sequential of sequential sequential	30 mmHg in chambers 50 to 30 mmHg gradient in chamber	N/A	Experimental study	bioimpedance, tank circumference, FASCO score, bioimpedance	Limb volume (water displacement)

B. SPECIAL SITUATIONS

Combined Lymphatic Surgery	Zahawi 1993	The reported primary objective of the study was to suggest for lymphedema in the limbs	IPC used before, during and after surgery for LVA and maintain suctions pressure, reduce wound complications and pain in excisional surgeries.	Higher pressure used for the more severe case 1, preoperatively.	3	secondary UE, LE	Case 1 (LE) combined with case 2 (LE) postop, case 2-30 mm Hg	Sequench	Lymphatics	case series	wound complications and pain in excisional surgeries.
C. NON-LIMBS LYMPHEDEMA											
Efficacy	Rasmussen 2019	Advanced pneumatic compression for treatment of lymphedema of the head and neck: a randomized, wait list controlled trial	Patients who received advanced pneumatic compression device (APC) case for lymphedema showed significant improvements in their perceived ability to control lymphedema, patient-reported outcomes, visible external swelling, and restricted pain compared to those who received wait list control.	significantly enhanced patient perception regarding their ability to control their lymphedema, potentially reducing patient distress.	43	secondary H&N	8w	23 to 45 minutes	7 days per week, but most are used once a day	N/A	Symptoms (including pain, heaviness, function (cervical motion, speech), swelling, internal and external)
Safety	Gonzalez 2019	Head and Neck Lymphedema: Response to Single and Multiple Sessions of Advanced Pneumatic Compression Therapy	The study evaluated lymphedema reduction after IPC completion finding a reduction in lymphedema in 6 out of 8 subjects and demonstrated improved facial composite measurements scores and a reported decrease in all 10 subjects with no adverse events reported.	The use of non-inflated, fluorescence lymphatic mapping (LFLM) by means of the superior of the pneumatic compression therapy, lymphatic drainage and the ability to observe enhanced lymphatic vessels and drainage, as well as changes in areas of dermal backflow, in subjects after the treatment.	10	secondary H&N	2 weeks	32m	7 days per week	N/A	external swelling (measured by LFLM), patient-reported symptoms, and subjective improvements
Imaging for Efficacy	Rasmussen 2019	Imaging the Lymphatic Response to Manual Lymphatic Drainage in Patients with Head and Neck Cancer-Related Lymphedema	Both MLD and IPC showed improvement in lymphatic velocity, and enhanced lymphatic drainage. The use of IPC showed a more comfortable, effective, and well-tolerated treatment with one treatment showing substantially significant reduction in overall symptoms but the reduction in trunk circumference.	The only study that used imaging after a sustained use of IPC	69	secondary H&N, limbs	2 weeks	N/A	daily for 2 weeks	N/A	ICGL before and 2 weeks after w. during IPC/MLD
Best Practice	Manowitz 2018	Advanced pneumatic compression to treat cancer-related head and neck lymphedema	Significant reduction in overall symptoms but the reduction in trunk circumference.	The best head use found to be safe, easy to use, and well-tolerated showing edema reduction after just a single initial treatment 100% compliance with IPC, Flat touch, used a study period long short	44	secondary H&N	Single session	32m	N/A	N/A	Patients reported comfort feeling post-treatment, and likelihood of home use, trunk circumference
Efficacy	Rahbar 2010	Advanced pneumatic compression for the treatment of lymphatic dysfunction in the upper limb	Re-allowing movement of edema fluid and improved lymphatic drainage and mobility. The study examines the change in limb volume and circumference after 2 hours of compression pressure to reduce chronic edema of the extremities	Determined sleeve pressure was required for patients to tolerate use of the device. Issues which can be associated with use of the device were noted. Indirect estimate of required compression pressure	12	secondary Trunk	10 days	1h	7 days per week	N/A	Patient reported symptoms, trunk circumference
D. SUPPORTIVE STUDIES ON COMPRESSION PRESSURE											
Efficacy	Zahawi 2017	Indo-cyanine green lymphatic mapping for evaluation of lymphatic function			N/A	UE, LE	single session			N/A	ICGL before and after, also measured limb volume using four different
Other	Funkh 2011	Dose finding for an optimal compression pressure to reduce chronic edema of the extremities			36 arm patients and 42 leg patients	UE, LE	N/A	N/A	N/A	N/A	Limb volume and circumference before and after treatment

ADL, activities of daily living; AE, adverse events; APC, advanced pneumatic compression device; ASE, Anson's Shoulder and Elbow Surgeons; BCR, breast cancer-related lymphedema; CB, compression bandages; CDT, complex decongestive therapy; CL, central vein; FU, follow up; ggs, groups; h/hr, hour; H&N, head and neck; ICG, indocyanine green; non-inflated lymphography; IPC, inflatable pneumatic compression; Jts, joints; LE, lower extremity; LGS, leg lymphedema complexity score; LYMAPCOL, lymphedema quality of life score; LSG, lymphoscintigraphy; LV, limb volume; LVA, lymphatic venous anastomosis; m, minute; MLD, manual lymphatic drainage; MOK, mechanism of action; NIRFLI, near-infrared fluorescence lymphatic imaging; NP-PCD, non-pneumatic pressure; non-compression device; PC-1, pneumatic compression device; PC-1, pneumatic compression device; PC-1, pneumatic compression device; PC-2, pneumatic compression device; PC-3, pneumatic compression device; PC-4, pneumatic compression device; PC-5, pneumatic compression device; PC-6, pneumatic compression device; PC-7, pneumatic compression device; PC-8, pneumatic compression device; PC-9, pneumatic compression device; PC-10, pneumatic compression device; PC-11, pneumatic compression device; PC-12, pneumatic compression device; PC-13, pneumatic compression device; PC-14, pneumatic compression device; PC-15, pneumatic compression device; PC-16, pneumatic compression device; PC-17, pneumatic compression device; PC-18, pneumatic compression device; PC-19, pneumatic compression device; PC-20, pneumatic compression device; PC-21, pneumatic compression device; PC-22, pneumatic compression device; PC-23, pneumatic compression device; PC-24, pneumatic compression device; PC-25, pneumatic compression device; PC-26, pneumatic compression device; PC-27, pneumatic compression device; PC-28, pneumatic compression device; PC-29, pneumatic compression device; PC-30, pneumatic compression device; PC-31, pneumatic compression device; PC-32, pneumatic compression device; PC-33, pneumatic compression device; PC-34, pneumatic compression device; PC-35, pneumatic compression device; PC-36, pneumatic compression device; PC-37, pneumatic compression device; PC-38, pneumatic compression device; PC-39, pneumatic compression device; PC-40, pneumatic compression device; PC-41, pneumatic compression device; PC-42, pneumatic compression device; PC-43, pneumatic compression device; PC-44, pneumatic compression device; PC-45, pneumatic compression device; PC-46, pneumatic compression device; PC-47, pneumatic compression device; PC-48, pneumatic compression device; PC-49, pneumatic compression device; PC-50, pneumatic compression device; PC-51, pneumatic compression device; PC-52, pneumatic compression device; PC-53, pneumatic compression device; PC-54, pneumatic compression device; PC-55, pneumatic compression device; PC-56, pneumatic compression device; PC-57, pneumatic compression device; PC-58, pneumatic compression device; PC-59, pneumatic compression device; PC-60, pneumatic compression device; PC-61, pneumatic compression device; PC-62, pneumatic compression device; PC-63, pneumatic compression device; PC-64, pneumatic compression device; PC-65, pneumatic compression device; PC-66, pneumatic compression device; PC-67, pneumatic compression device; PC-68, pneumatic compression device; PC-69, pneumatic compression device; PC-70, pneumatic compression device; PC-71, pneumatic compression device; PC-72, pneumatic compression device; PC-73, pneumatic compression device; PC-74, pneumatic compression device; PC-75, pneumatic compression device; PC-76, pneumatic compression device; PC-77, pneumatic compression device; PC-78, pneumatic compression device; PC-79, pneumatic compression device; PC-80, pneumatic compression device; PC-81, pneumatic compression device; PC-82, pneumatic compression device; PC-83, pneumatic compression device; PC-84, pneumatic compression device; PC-85, pneumatic compression device; PC-86, pneumatic compression device; PC-87, pneumatic compression device; PC-88, pneumatic compression device; PC-89, pneumatic compression device; PC-90, pneumatic compression device; PC-91, pneumatic compression device; PC-92, pneumatic compression device; PC-93, pneumatic compression device; PC-94, pneumatic compression device; PC-95, pneumatic compression device; PC-96, pneumatic compression device; PC-97, pneumatic compression device; PC-98, pneumatic compression device; PC-99, pneumatic compression device; PC-100, pneumatic compression device.

upper limb (1 study, acquired disease) lymphedema on a sample size ranging from 12 to 1065 patients (1,10,13,20-24). The treatment sessions varied from 45 minutes to 1 hour/day for 5-7 days a week and patients continued to use compression garments in between sessions. Peak IPC pressures used were 80 to 120 mm Hg in lower limbs and 60 mm Hg in the upper limb. In one study, 74 patients experienced significant reductions in ankle and calf circumference, a decrease in cellulitis rates from 32% to 12%, and enhanced quality of life scores after 12 weeks of daily 1-hour IPC sessions, with the benefits sustained at 1 year (20). After at least 3 months of IPC treatment all patients experienced symptoms improvement and 54% experiencing substantial quality of life enhancements and significant reductions in limb girth and infection episodes (21).

Prolonged use of IPC (more than 8 weeks) was associated with improved quality of life, limb volume reduction, and enhanced limb function in two other studies (1,21). Improved tissue elasticity besides significant reductions in limb circumference was observed in 18 patients with lower limb lymphedema over a three-year period (13). Improved skin fibrosis and reduced limb pain and was reported in majority of the participants with lower extremity lymphedema in another study, although only a third of the patients showed >10% volume reduction (10).

The only study that exclusively studied utility of IPC without MLD in upper limbs randomized the postmastectomy arm lymphedema patients into study and control groups but used shorter treatment period. After two cycles of 2 hours per day of 2 weeks treatment, the mean volume change was significantly more in the IPC group ($p=0.009$) compared to the control ($p=0.3$), although no difference in the number of patients who achieved > 20% reduction was observed (22-25).

These studies collectively indicate that IPC when used over several weeks can be effective as a standalone therapy for managing lower extremity lymphedema. However, the undisclosed variation in the duration and strength of the compression garments is a potential confounder. While volume reduction

was frequently noted as significant, only a few studies reported precise percentage reductions. Additionally, the lack of a control group in most of the above studies limits the interpretation of results. Due to paucity of studies, effectiveness of IPC alone for managing arms lymphedema cannot be determined.

IPC vs MLD

In the comparative analysis between IPC and MLD, four studies examined treatments for upper (3 studies) or both upper and lower limbs (1 study) (26-29). The duration of IPC treatments ranged from 30 minutes to 2 hours for 3 -5 days a week, using compression pressures of 25 mm Hg to 80 mm Hg. These investigations reported that IPC and MLD resulted in similar decreases in limb volume as well as improvement in pain levels, range of motion, and patient-reported symptoms over the study period of 2-6 weeks in 28 to 182 subjects (27). Volume reduction was more pronounced in the lower limbs as compared to arms using either therapy. Self-lymphatic drainage was permitted along with IPC in one study making the findings less reliable (27). Overall, the results of rest of the included studies indicated that both treatment approaches yielded comparable outcomes in terms of volume reduction and symptomatic relief, although the time and cost consumption for MLD was higher and compliance was lower (28).

IPC and MLD vs. MLD

Seven prospective studies examined IPC as an adjunct to MLD, of which six studies focused on arms lymphedema and only one studied lower limbs. In the context of lower extremity lymphedema, two studies indicated that combining high-pressure IPC (120 mm Hg) with MLD led to greater reductions in limb volume compared to MLD alone (30,31). Furthermore, incorporation of IPC into CDT was associated with improved compliance and significantly lower rates of infection (16.9% vs 29.2%), hospital admissions (13.6% vs 29.2%), and fewer physical therapy sessions over 18 months among 62 patients (30). In contrast,

six studies investigating upper extremity lymphedema produced variable results (31-36). Two studies found that combining MLD + IPC at 40-60 mmHg pressure for 30-60 minutes daily for 2 to 52 weeks, resulted in superior volume reduction and/or limb function improvement compared to MLD without IPC (32,33). However, the remaining four studies did not observe significant benefits which could be related to lower pressure levels (30-40 mmHg), shorter treatment durations (30 minutes), or limited sample size in these studies (31,34-36).

IPC as an adjunct to Surgical Treatment

One study investigated the perioperative application of IPC to improve outcomes of lymphatic surgery (37). The study involved 15 patients undergoing lymphaticovenular anastomosis, during which the use of IPC enhanced the visualization of lymphatic vessels intraoperatively. The IPC was continued postoperatively to promote flow through the anastomoses. When used prior to excisional lymphatic surgeries, authors observed a decrease in wound complications and a reduction in postoperative pain. These findings suggest the potential of utilizing IPC for improving lymphatic surgery outcomes and further research may be steered in this direction.

Application in Head and Neck

Four studies evaluated use of IPC for management of head and neck lymphedema (19,38-40). Mayrovitz et al reported high levels of patient satisfaction and notable reductions in facial and neck lymphedema following a single session of IPC lasting 32 minutes (38). Gutierrez et al documented a decrease in dermal backflow as shown by indocyanine green lymphography in 6 out of 8 patients, following two weeks of daily treatment sessions of the same duration (39,41). Ridner et al and Rasmussen et al further corroborated these outcomes, with patients reporting swelling reduction, improved symptoms and enhanced quality of life following 2 to 3 months of daily IPC application (19,40).

Application in Trunk and Chest

Only one study was found that specifically evaluated role of IPC in managing trunk lymphedema. All 12 patients reported improved symptoms but there was no significant reduction in the trunk volume after 10 days of 1-hour daily IPC treatment (42).

Factors Influencing the Effectiveness of IPC Treatment

In a cohort of 56 patients, Forner-Cordero et al found that similar to other decongestive therapies, lower limb lymphedema and less advanced disease showed more notable improvements with IPC as compared to upper limbs and advanced lymphedema (28). Conversely, in a larger cohort of 208 patients, Muluk et al did not find an inverse relationship between disease severity and volume reduction (10). This conflicting evidence regarding the correlation between disease severity and response to IPC mirrors findings reported for CDT (43,44). High BMI, which has been linked to poorer responses to CDT, has yet to be thoroughly evaluated for its impact on IPC outcomes.

Durability of Treatment Results

Sixteen studies investigated the longevity of results post IPC treatment, with follow-up periods extending up to one year. It was generally noted that volume reductions achieved with IPC require ongoing use of compression garments. The outcomes tend to diminish once IPC is discontinued (11,45). For example, Johansson et al reported a 31% reduction in lower limb volume at the conclusion of the IPC treatment period, which declined to a 25% reduction after one month of stopping the treatment (26). These findings are similar to other decongestive therapies and underscore the importance of continued self-care with IPC for maintaining lymphedema reduction (11,26,32,46).

Machine Settings

Pressure Setting for Lower Extremity

In their investigation to determine adequate compression, Olszewski et al and Zaleska et al conducted tissue pressure measurements during IPC treatments (13,47-52). Effective therapeutic subcutaneous pressures greater than 40 mmHg and improved proximal fluid movement on lymphoscintigraphy were achieved with IPC machine pressures of 80–120 mmHg for durations exceeding 60 seconds (11,16). The authors concluded that higher interface pressure is necessary as the severity of lymphedema increases due to greater tissue stiffness (48,50). Furthermore, using a combination of ICG lymphography and tissue pressure measurements, Zaleska et al found that IPC pressures below 80 mmHg were insufficient for lymph clearance in advanced stages of the disease. Supporting these findings, Taradaj et al demonstrated in a clinical study that limb volume reduction in patients treated with IPC at 60 mmHg did not statistically differ from the no-IPC group but was significantly less than in the group treated with higher pressure (120 mmHg) (31).

Pressure Setting for Upper Extremity

Bok et al conducted a comparison of IPC pressures of 25, 35, and 45 mmHg in a cohort of 40 patients (53). Utilizing ultrasound and acoustic radiation force impulse imaging, they assessed change in tissue thickness and stiffness with IPC. The findings indicated that the 35-mmHg pressure setting was optimal, as it provided significant volume reduction while maintaining a balance between patient comfort and treatment efficacy.

Compression Cycle

Most current IPC machines are designed so that the distal chamber remains inflated while the other chambers sequentially inflate and deflate during the compression cycle (50). Zaleska found that to generate effective tissue pressures for lymphatic outflow, high-pressure cuffs should remain inflated for more than 50 seconds (50). Additionally, Pilch et al studied

the effects of different inflation/deflation cycles (60/60 seconds versus 90/30 seconds) in a cohort of 81 patients and found no significant differences in outcomes between the two settings (54). One study evaluated the benefit of MLD mimicking truncal compression prior to limb compression as compared limb pneumatic compression alone and did not find significant difference in the outcomes (55).

Treatment Duration

The treatment periods used in the published studies have ranged from 30 mins to 2 hours per day for 3 to 7 days per week. Keeley et al randomized 21 patients to receive 1 hour of IPC once daily, twice daily, or 2 hours twice daily over a span of 5 days (56). The study concluded that extending IPC beyond 1 hour daily does not yield additional reduction in limb volume or tissue extracellular fluid percentage.

Cost Effectiveness

The economic impact of IPC was explored in six studies. Lerman et al benchmarked IPC against other compression therapies in a group of 138 patients, identifying significant cost savings over 18 months attributed to fewer complications and hospital stays (57). IPC users experienced a reduction of 66% in emergency department visits and 69% in hospitalizations. Desai et al reported substantial savings, amounting to \$3,200 per patient annually (23). Meanwhile, Karaca-Mandic et al presented IPC as a cost-effective solution when considering quality-adjusted life years, with an incremental cost-effectiveness of \$1,400 (58). The amount of cost-benefit of IPC devices varied depending on device pricing and the extent of insurance coverage.

DISCUSSION

The evolution of Intermittent Pneumatic Compression (IPC) technology provides a therapeutic opportunity for personalized, patient-centered care. As lymphedema remains a chronic, often debilitating condition, the

diversification of IPC devices – with their varied pressure settings, sleeve designs, and compression modes – reflects a broader trend of tailoring treatment to individual patient needs. Our review illuminates the complex landscape of IPC, where the multitude of device configurations and operational parameters mirrors the heterogeneity of lymphedema presentations and patient experiences. Despite the availability of numerous studies, lack of standardized protocols and conflicting evidence continues to obscure the optimal application of IPC, necessitating further research to unify these disparate threads into a coherent clinical strategy.

It is important to note that although many studies compared outcomes of IPC with reference to MLD, the efficacy of MLD as part of CDT itself has remained questionable (59). Central to the value of IPC is its capacity to empower patients, offering a semblance of control over their condition. This autonomy is especially critical given the limitations in access to specialized lymphedema services and trained therapists. The convenience and adaptability of IPC facilitate consistent self-management, potentially enhancing adherence to therapeutic regimens and improving long-term outcomes.

The efficacy of IPC, however, is not universally consistent across different patient populations and lymphedema manifestations. Our review suggests a greater benefit in lower extremity lymphedema, possibly due to greater struggle for patients to mobilize the lymphedema fluid against gravity in this region. The nuanced interplay between IPC treatment parameters and patient-specific factors – such as disease severity, and body mass index – highlights the importance of individualized treatment protocols. The optimal IPC settings, notably pressure and duration, should be tailored based on comprehensive assessments including imaging studies and tissue pressure measurements, to maximize therapeutic efficacy and patient comfort. An interesting trend shows that recommended pressure settings have decreased over time, with European studies typically using higher pressures than U.S.

studies – possibly because of difference in the disease severity. The presence of lipodystrophy and fibrosis would require higher pressure settings for efficacy, which although would increase patient discomfort and reduce compliance. While there are several studies evaluating pressure settings for leg lymphedema (which generally recommend high pressure), there is limited research on optimal pressure settings for arm lymphedema.

From a mechanistic perspective, IPC acts on both lymphatic and interstitial fluid dynamics, yet the relative contributions of these pathways remain poorly understood (15,48). Imaging studies during IPC sessions have shown enhanced lymphatic propulsion, suggesting a direct effect on the lymphatic system. However, the long-term implications of these acute changes, particularly regarding the sustainability of lymphatic function improvement, are yet to be fully explored. This gap underscores the need for longitudinal research to elucidate the mechanisms underlying IPC's therapeutic benefits and to define its role within the broader spectrum of lymphedema management strategies.

The application of IPC extends beyond routine outpatient care, showing potential in perioperative settings to enhance surgical outcomes in lymphedema treatment. Preliminary evidence indicates that IPC can improve lymphatic vessel visualization during surgery, reduce postoperative complications, and facilitate quicker recovery. Additionally, emerging studies highlight the utility of IPC in managing truncal and head/neck lymphedema, areas that have historically received less attention in lymphedema research. These findings point to a promising avenue for expanding IPC's clinical utility, warranting more comprehensive investigations to validate these initial observations and refine treatment protocols.

CONCLUSION

IPC emerges as a safe and economical option for management of extremities' lymphedema, aligning well with patient-led care approaches. Future research should aim to refine IPC protocols and ascertain its long-

term benefits and its integration into contemporary lymphedema treatment regimes.

CONFLICT OF INTEREST AND DISCLOSURE

All authors declare that no competing financial interests exist in products or devices mentioned in the manuscript.

AUTHOR CONTRIBUTIONS

Sonia Pandey coordinated the study and composed the original manuscript. Berk Ozmen, Suat Morkuzu, Ying Xiong, Elise Kemp assisted with the articles screening and manuscript drafting. Wei F. Chen conceptualized the project and edited the manuscript.

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TABLE 1
Detailed Manuscript Information Grouped into 4 Groups used for Analysis
(Limb Lymphedema, Special Situations, Non-Limb Lymphedema,
and Relevant Compression Pressure Studies)

Aim	Research focus	Author, Year	Title	Summary of Outcomes	Salient point	Sample size	Etiology	Region	Total Duration of Use	Duration per Day	Max Follow up period	Frequency	Compression Cycles	IPC Pressure	Device name	Study Design	Outcome measures	Adverse events	
A. LIMB LYMPHEDEMA																			
IPC without MLD	Efficacy	Frejne 2022	Severe lymphoedema in gynaecological cancers:	Following IPC participants showed a significant improvement in mean global LYMQOL scores, physical component of Superior Clinical, Quality of Life, Functional, and Health	IPC improves all patients' functional and symptom scores, except sexual function	12	secondary	LE	8w	1h	N/A	daily	N/A	100 mmHg	N/A	Prospective cohort study	Quality of life scores and history of hospital		
	Efficacy	Maldonado 2021	Assessment of quality of life changes in patients with	IPC treatment significantly improved LYMQOL scores, physical component of Superior Clinical, Quality of Life, Functional, and Health	The Flexitouch APCD showed a significant limb girth reduction as early as 12 weeks	74	secondary, primary	LE	52w	45m	52 weeks	5 days per week	N/A	N/A	Flexitouch	Prospective cohort study	The primary endpoints: QOL change at 12 weeks		
	Efficacy, Cost	Desai 2019	Superior Clinical, Quality of Life, Functional, and Health	IPC resulted in a 28% decrease in absolute limb volume, decreased body mass index	IPC improves clinical outcomes, quality of life, and function for UE and LE secondary lymphedema	117	secondary	UE, LE	52w	1h			twice daily	18 s per chamber	40 or 50 mmHg	Metrum Cryoflex	Prospective cohort study	Absolute limb volume, BMI, SF-36 quality of life, and leg	
	Safety, Efficacy	Blumberg 2016	Pneumatic Compression Improves Quality of Life in	The study showed a significant decrease in ankle and calf limb girth, as well as the	IPC improves symptoms of LE lymphedema	100	secondary, primary	LE	57w	45m			5 to 7 days per week	N/A	N/A	N/A	Prospective cohort study	Episodes of cellulitis, number of ulcers, venous	
	Efficacy, Cost	Brayton 2014	Lymphedema prevalence and treatment benefits in cancer: impact of a	The use of IPC was associated with a significant decrease in rates of hospitalizations, outpatient hospital visits,	Lymphedema is common among cancer survivors, and IPC is valuable in treating lymphedema	1065	secondary	N/A	N/A	N/A			N/A	N/A	N/A	N/A	Retrospective cohort study	Rates of hospitalizations, outpatient hospital visits, cellulitis diagnoses, and	
	Safety, Efficacy, Settings	Zaleska 2014	The Effectiveness of Intermittent Pneumatic Compression	The limb circumference decreased or did not further increase, elasticity of tissues increased or was maintained. No	The first one hour IPC showed decrease in circumference of lower parts of the calf and thigh with increase below knee and in the	18	secondary, primary	LE	2 to 3 years	45 mins					80 to 120 mmHg	N/A	prospective cohort study	limb circumference, tissue Tonometry	Transient increase in groin and knee swelling, although calf and thigh reduced. No
	Efficacy, Safety	Muluk 2013	Pneumatic compression device for management of lower limb lymphedema	In patients treated with APCD, 90% experienced a significant reduction in limb	The APCD treatment was associated with significant reductions in LV and showed	196	secondary, primary	LE	N/A	N/A			4 to 9 weeks	N/A	80 to 110 mmHg	N/A	Prospective cohort study	Reduction in limb volume (LV), clinician and patient	Four adverse events were recorded for these subjects
Efficacy	Modagheh 2010	A newly designed IPC device for management of lymphoedema	IPC at high pressure was effective in reducing affected lower limb edema by a mean of 75% in lymphedema patients,	Introduction of a new mode for IPC which allows unidirectional forward lymph flow and is well-tolerated in chronic cases of the	43	secondary, primary	LE	2 days	8h			single in hospital treatment	N/A	80 to 120 mmHg	N/A	Experimental study, single session	Limb circumference		
Safety	Borris 1998	The risk of genital edema after external pump	Of the 128 patients with lower limb lymphedema, 75 received no pump	IPC can increase proximal swelling, no mention on the duration for which the	128	secondary, primary	LE								N/A			Genital swelling	
IPC vs MLD	Efficacy	Gozza 1996	Pneumatic compression vs	After 9 weeks of treatment, mean delta	IPC may have limited clinical value	67	secondary	UE	9w	2h			N/A	60 mmHg	N/A	RCT	Limb circumference before		
	Efficacy	Sanal-Toprak 2019	The efficacy of intermittent pneumatic compression as a	No significant difference between the two groups in the extent of improvement in	IPC treatment can be substituted for MLD in CDT due to its easier accessibility and	46	secondary	UE	5w	30 mins	3 months	3 days per week	Sequential	50-80 mmHg	N/A	RCT	circumference, shoulder pain and ROM		
	Efficacy	Gurdal 2012	Comparison of intermittent pneumatic compression with	Both treatment modalities, namely IPC + Self Lymphatic Drainage (SLD) and	Manual lymphatic drainage and compression bandages may perform	30	secondary	UE	6w	45m			3 days per week	N/A	25 mmHg	N/A	NRCT	Limb circumference before and at the 1st, 3rd, and 6th	
	Efficacy	Fomer-Cordero 2021	Physical therapy in the decongestive treatment of lymphedema: A	Group A (control group): manual lymphatic drainage + Intermittent Pneumatic Compression +	IPC and bandages is not inferior to IPC + manual lymphatic drainage + bandages	182	secondary, primary	UE, LE	4w	30m	2 months	5 days per week	N/A	50 to 80 mmHg	N/A	RCT	Limb volume before and after treatment	discomfort and lymphangitis	
	Efficacy	Johansson 1998	A randomized study comparing manual lymph	Arm volume, mobility, strength and symptoms improvement not sig different in	IPC is as efficacious as MLD in BCRL	28	secondary	UE	2w	2 hrs			5 days per week	40-60 mmHg	N/A	RCT	arm volume, level of mobility and strength		
IPC + MLD	Efficacy	Soran 2022	Adding Pneumatic Compression Therapy in	IPC significantly reduces infection rates, hospital admissions, and physical therapy	Compliance improved with IPC	69	secondary, primary	LE	78w				Sequential	N/A	N/A	NRCT	Rate of infection, number of hospital admissions, and		
	Efficacy	Szolonsky 2022	IPC acts synergistically with MLD in complex decongestive physiotherapy	Arm volume and symptoms significantly improved in both groups at therapy end, at 1 month and at 2 months. Volume reduction	Excellent discussion about fears of using IPC and the different pressures.	27 (13 CDT, 14 CDT+IPC)	secondary	UE	2w	60 mins MLD vs 30 mins MLD + 30 mins IPC	2 months	5 days/Wk		50 mmHg	Lymphamat	RCT	Limb volume, symptoms questionnaire		
	Efficacy	Tastaban 2020	Role of intermittent pneumatic compression in the treatment of breast	IPC seems to add no benefit when combined with CDT for lymphedema, but, may be functional in reducing the		76 (38 CDT, 38 CDT+IPC)	secondary	UE	4w	60 mins MLD vs 60 mins MLD + 30 mins IPC			5 days per week		30-40 mmHg	Pulse press	RCT	Percentage reduction of excess volume (PREV), symptoms of pain,	
	Efficacy	Uzkeer 2015	Efficacy of manual lymphatic drainage and	All tested parameters of limb volume, circumference and dermal thickness	no additional benefit of adding IPC to CDT for BCRL	31	secondary	UE	3w	45 mins			6 days per week		40 mmHg	N/A	RCT	Limb circumference measure means, volume	
	Efficacy	Uzkeer 2013	Intermittent pneumatic compression pump in upper	Significant improvements were observed in the function (ADL) and ROM of the	no additional benefit of adding IPC to CDT for BCRL	25	secondary	UE	3w				5 days per week		60 mmHg	N/A	RCT	ROM of UE JTs	
	Efficacy	Haghighat 2010	Comparing two treatment methods for post	The study showed that both the use of CDT alone and in combination with IPC	CDT alone provided better results in reducing limb volume than when combined	112	secondary	UE	12w	1h	12w	5 days per week	N/A	40 mmHg	N/A	RCT	The volume reduction of the upper limb measured by		
	Safety, Efficacy	Szuba 2002	Decongestive lymphatic therapy for patients with	The addition of IPC to standard therapy led to a significant increase in mean volume		23	secondary	UE	26 - 52 weeks	30 to 60 minutes			N/A	Sequential	40 to 50 mmHg	N/A	RCT	Volume - water displacement	No adverse effects on skin elasticity and joint range of
Pressures	Pressures	Taradaj 2015	Comparison of efficacy of the intermittent pneumatic	IPC at 120 mmHg led to the greatest reduction of edema compared to lower		81	Phlebolymph edema	LE	4w	45m			Sequential	60 or 120 mmHg	N/A	NRCT			
	Frequency of use	Keeley 2023	A Prospective Preliminary Study Examining the Physiological Impact of Pneumatic Compression Dosing in the Treatment of Lower Extremity Lymphedema	Limb volume measurements demonstrated no potential benefit of increasing IPC duration more than 1 hour per day during 5 days of treatment	No benefit of > 1 hr per day IPC in short FU	21	primary-secondary	LE	5 days or 12 days	Group A: 1 hour. Group B: 1 hour (but twice per day, so 2 hours total). Group C: 2 hours (but twice per day, so 4 hours total).	5 days	Group A: Once per day Group B: Twice per day Group C: Twice per day	N/A	55 mmHg distal to 30 mmHg proximal	N/A	Prospective cohort study	Changes in limb volume (LV), tissue fluid, tissue tone, and patient-reported outcomes (PROs) -> only volume and BIS were reliable		
	Compression modes	Dunn 2022	Intermittent Pneumatic Compression for the Home-Based Intermittent Pneumatic Compression	The study found that the LymphAssist group exhibited no significant volume	The improvement in limb volume and quality of life for patients with LE	40	secondary, primary	LE	5w	N/A			twice daily	N/A	40 mmHg	LymphAssist	RCT	Bilateral leg volume (with circumference) and quality	
	Compression modes	Kim 2022	The use of a home-based Intermittent Pneumatic Compression Therapy: The Impact in Chronic Leg Lymphedema in Patients Treated for Gynecologic Cancer	The use of a home-based IPC device alongside a routine self-maintenance program of short-stretch bandages in stage 3 chronic leg lymphedema patients resulted in a lower limb-volume difference ratio and higher quality of life compared to using a home-based IPC device alone.	The IPC device has a distinct mode designed to imitate the manual lymphatic drainage (MLD) technique, aiming to gently aid lymphatic draining of the proximal extremities. This study shows that using such an IPC device with an MLD-mimicking program can be beneficial in maintaining limb volume and enhancing the QOL for patients with stage 3 chronic leg lymphedema during their maintenance phase.	30	secondary	LE	N/A	1h	4 weeks	twice a day for 4 weeks	MLD-mimicking mode: 3 s inflation, 1 s holding time, 7 s deflation and resting time; conventional mode: 6 s inflation, 1 s holding time, 7 s deflation time for each chamber	MLD-mimicking mode: 40-60 mmHg; conventional mode: 80-100 mmHg	N/A	NRCT; Prospective cohort study	Limb-volume measurement, quality of life (QOL), satisfaction, and safety of the IPC device.		
	Programmability	Karac-Mandic 2017	A comparison of programmable and non-programmable	Dynamic pressure programmable devices demonstrated better lymphedema-related health outcomes compared to	The study compared the clinical and health utilization outcomes between two types of PCDs (NP-PCD and P-PCD) in patients	1731	secondary, primary	unspecified	12 months	N/A		12 months	N/A	N/A	N/A	N/A	Retrospective cohort study	Rates of lymphedema-related cellulitis, manual therapy use, outpatient	

Device Settings and modes - Clinical studies	Comparing 2 devices	Fife 2012	A randomized controlled trial comparing two types of pneumatic compression for breast cancer-related lymphedema treatment in the home	The APCD-treated group showed a significant 29% reduction in edema compared to a 16% increase in the IPC group, with a 53% reduction in mean TDC values for the APCD group and a 19% increase for the IPC group, indicating that adjuvant treatment with an APCD provides better outcomes than with an IPC, and both groups demonstrated very good device compliance.	Advanced IPC as an adjuvant treatment may be better than standard IPC devices for treating BCRL. Seven pts had adverse events, more in the Std gp.	36	secondary	UE	12w	1h		N/A	18 s per chamber (std) sequentially inflated ending with all in inflated vs 1 to 3 secs per chamber (only one inflated at a time)	30 mmHg (standard) or n 9 04: 4.2 mmHg and 13.7±4.8 mmHg (Advanced)	N/A	RCT	Limb volume (with circumference) and tissue water measurement (with Moisture meter) at Week 0 and 12	Seven patients had adverse events, more in the standard group.	
	Trunk compression preceding arm compression	Rahner 2012	A randomized clinical trial comparing advanced pneumatic truncal, chest, and arm treatment to arm treatment only in self-care of	When comparing experimental truncal chest/arm advanced pneumatic compression therapy to arm-only pneumatic compression (control), there were no statistically significant changes in		42	secondary	UE	30 days	1h in UE + trunk gp, 36 mins in UE only gp		7 days per week	N/A	9.0 ± 4.2–13.7 ± 4.9 mm Hg	N/A	RCT	LSIDS-A score, arm circumference, trunk circumference, (FASQ score, bioimpedance)		
	No. of chambers and cycle time	Pulch 2009	Influence of compression cycle time and number of sleeve chambers on upper extremity lymphedema	IPC is an effective method of volume reduction in women with postmastectomy arm lymphedema regardless of cycle times and number of sleeve chambers. Two			57	secondary	UE	5w	1h		5 days per week for 5 weeks	N/A	30 to 50 mmHg	N/A	RCT	Limb volume	
	Comparing 2 devices	Maynovitz 2007	Interface pressures produced by two different types of lymphedema therapy devices	Significant differences in pressure timings, patterns, and magnitude were found after comparing the Lympha Press and Flexitouch devices. In terms of timing, the duration of pressure pulses in the Flexitouch device were significantly shorter than that of the Lympha Press. Due to this quick application of pressure, the	There were significant differences between the Flexitouch and Lympha Press devices. During inflation cycles, the pressures exerted by the Lympha Press were higher and were sustained for longer periods of time. However, the Flexitouch differed drastically by displaying quick rise and fall progressions		10	N/A	UE	Single session for each device	Single session for each device	N/A	Single session for each device	Sequential	45mmHg for the Lympha Press and the "standard" setting for Flexitouch	Lympha Press, Flexitouch	Experimental study	Interface pressures applied to left forearm (used 256-pressure sensor array) over 0.1 second intervals for at least 2 cycles of each device	
	No. of chambers	Bergan 1998	A comparison of compression pumps in the treatment of lymphedema	One and three chamber garments were less effective than 10 chamber garment in reducing swelling	multi chamber sequential compression more effective than fewer chambers		35	secondary, primary	UE, LE	N/A	2 hrs		N/A	Sequential or non sequential	50 mmHg in one or 3 chambers, 80 to 30 mmHg gradient in ten chambers.			Limb volume (water displacement)	
Experimental Studies	Imaging for Efficacy, Best settings	Zaleska 2019	The Effectiveness of Intermittent Pneumatic Compression in Lymphedema of Lower Limbs: Methods of Evaluation and Results	Adjustment of compression parameters to tissue stiffness, fluid accumulation volumes, and fluid movement ability (hydraulic conductivity of tissues) at various limb levels is indispensable for effective therapy. The recommended differential compression pressures and prolonged timings at various limb levels		52	secondary, primary	LE	1 hr				5 sec inflation, 5 sec deflation for each chamber, total 40 sec period for 1 cycle.	The sleeve inflation pressure at foot level was 120 mmHg, gradually decreasing in the groin by 20%.	N/A	Experimental study, single session	tissue stiffness, fluid pressure, flow volume, LSG, ICGL		
	Efficacy, Best settings	Bok 2018	Evaluation of Stiffness in Postmastectomy Lymphedema Using	After a single session of IPC, using a pressure of 35 mmHg resulted in the largest improvement in proximal upper arm	IPC reduces stiffness and subcutaneous tissue thickness of UE in patients with BCRL	45	secondary	UE	single session	N/A		N/A	N/A	25, 35, and 45mmHg	N/A	Experimental study, single session	UE subcutaneous tissue thickness, circumference, and stiffness		
	Imaging for Efficacy	Aldrich 2017	Effect of pneumatic compression therapy on lymph movement in	The study demonstrate improved lymph movement during and after IPC in all affected legs of the subjects tested, with	IPC improves lymphatic function either by mobilizing fluid through lymph vessels or through interstitial channels.	4	secondary, primary	LE	single session	1h			N/A	45 distal to 30 mmHg proximal	N/A	Experimental study, single session	ICGL before (15 mins after injection), during and after IPC using compression		
	Imaging for Efficacy	Kitayama 2017	Real-Time Direct Evidence of the Superficial Lymphatic Drainage Effect of Intermittent Pneumatic	Different inflation/deflation modes and two different pressures evaluated (45 and 90 mmHg) using quantitative ICGL and software-assisted video analysis. ICG	Preop severity grading done with LSG and those with dermal backflow in leg and foot were excluded (only mild cases included).	25 (8 control, 17 affected limbs)	secondary	LE	single session	single session		single session	45 mmHg and 90 mm Hg	N/A	Experimental study, single session	ICGL- lymph velocity, real time with transparent compression sleeve, during IPC			
	Best settings	Zaleska 2013	Pressures and timing of intermittent pneumatic compression devices for	The study points to the necessity of applying high pressures (120 mmHg) at ankle with 20 % proximal pressure gradient	The pneumatic devices were set at three different inflation times 5, 20, or 50 sec in each chamber of the eight chambers, and	18	secondary, primary	LE	single session						N/A	Experimental study, single session	subcut pressure		
	Efficacy, MOA	Oliszewski 2011	Tissue fluid pressure and flow during pneumatic compression in lymphedema of lower limbs	The skin's rigidity (fibrosis) and the dissipation of applied compression force to proximal noncompressed limb regions resulted in a high pressure gradient through	Tissue fluid pressures generated by a pneumatic device were found lower than in the compression chambers. The obtained results point to the necessity of applying	15	N/A	LE	single session	single session		single session	sequential, distal most chamber was not deflated	50 to 120 mmHg	Biocompression	Experimental study, single session	Continuous Tissue pressure, and girth (plethysmograph attached to strain gauges applied at different limb)		
	Imaging for Efficacy	Oliszewski 2011	Pathways of lymph and tissue fluid flow during intermittent pneumatic	The study utilized LSG to demonstrate the movement of subcutaneous fluid and static lymph propelled towards the groin after	The study finds that while intermittent pneumatic compression can effectively push mobile tissue fluid towards the groin,	15	secondary	LE	single session	N/A			N/A	N/A	50 to 125 mmHg	N/A	Experimental study, single session	LSG- The primary outcomes are the pathways and destinations of fluid moved	One patient experienced muscle cramps and a second reported increased limb
	Imaging for Efficacy	Adams 2010	Direct evidence of lymphatic function improvement after advanced pneumatic	Improvement defined as proximal movement of dye after therapy. Lymphatic function improved in all control subjects	lymphatic function locally and systemically	3 controls, 6 with BCRL	secondary	UE	single session	single session. An hour-long IPC (30 mins trunk and proximal arm basins			single session		NA	Flexitouch	Experimental study, single session		
BI. SPECIAL SITUATIONS																			
Efficacy in fibrinosis	Manjula 2002	Evaluation of sequential intermittent pneumatic	12 patients with grade II fibilar lymphoedema had >26% reduction in			28	secondary	LE	4w			6 months						volume (with water displacement)	
Efficacy in children	Hassall 2001	A Retrospective Study of the Effects of Lympha Press Pump on Lymphedema in A	In this retrospective study of 16 children with lymphedema of the upper or lower extremities, the Lymphassess pump	pressure required decreased over time in the same pt		16	secondary, primary	UE, LE	N/A	N/A		N/A	N/A	N/A	Lympha Press	Retrospective cohort study	Volume and circumference of upper or lower limbs before and after treatment		

Combined with Lymphatic Surgery	Zelikovski 1983	The sequential pneumatic compression device in surgery for lymphedema in the limbs	IPC used before, during and after surgery. Aids in trap lymphatics visualization for LVA and maintains anastomosis patency, reduce wound complications and pain in excisional surgeries.	Higher pressure used for the more severe case 1, preoperatively.	3	secondary LE, UE	1 UE, 2 LE	Case 1 (LE)- 36 hrs continuously pre skin and subcut resection, 3 days postop 6 hrs/Day, Case 2 (LE)- introp for dilating groin lymphatics for LVA, postop for 12 hrs, Case 3 (UE) intraop for propelling blue dye, skin and subcut resection and LVA done.					Sequential	case 1 - 130 mm Hg preop, 80 mm Hg postop, case 2- 80 mm Hg	Lymphpress	case series	wound complications and pain in excisional surgeries.	
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C). NON LIMBS LYMPHEDEMA

Safety, Efficacy	Ridner 2021	Advanced pneumatic compression for treatment of lymphedema of the head and neck: a randomized wait list controlled trial	Patients who received advanced pneumatic compression device (AIPC) care for lymphedema showed significant improvements in their perceived ability to control lymphedema, patient-reported outcomes, visible external swelling, and reported pain compared to those who	significantly enhanced patient perception regarding their ability to control their lymphedema, potentially reducing patient distress.	43	secondary	H&N	8w	23 to 45 minutes			prescribed twice daily, 7 days per week, but most pts used once a day	N/A	N/A	N/A	RCT	Symptoms (including paresthesia, pain, heaviness), function (cervical motion, speech), swelling in internal and external	
Safety, Efficacy	Gutierrez 2019	Head and Neck Lymphedema: Treatment Response to Single and Multiple Sessions of Advanced Pneumatic Compression Therapy	The study evaluated ICGL-dermal backflow area in 10 patients before and after IPC completion, finding a reduction in backflow in 6 out of 8 subjects, and demonstrated improved facial composite measurements scores and self-reported outcomes in all 10 subjects with no adverse events reported.	The use of near-infrared fluorescence lymphatic imaging (NIRFLI) to visually assess the impact of the pneumatic compression therapy on lymphatic drainage and the ability to observe enhanced lymphatic uptake and drainage, as well as changes in areas of dermal backflow, in subjects after the treatment.	10	secondary	H&N	2 weeks	32m	2 weeks	7 days per week	N/A	N/A	Flexitouch	Prospective cohort- Observational study where subjects were imaged using NIRFLI before and after the initial treatment, and then again after 2 weeks of daily treatment.	external swelling (measured using facial composite scores), and subject-reported improvements.	No adverse events were reported	
Imaging for Efficacy	Rasmussen 2019	Imaging the Lymphatic Response to Manual Lymphatic Drainage and	Both MLD and IPC showed improvement in lymph velocity, and enhanced lymphatic uptake in both control and symptomatic	The only study that used imaging after a sustained use of IPC	69	secondary	H&N, limbs	2 weeks	N/A		daily for 2 weeks	N/A	N/A	N/A	NRCT- conference abstract	ICGL before and 2 weeks after +/- during IPC/MLD		
Safety, Efficacy, Best Settings	Mayrovitz 2018	Usability of advanced pneumatic compression to treat cancer-related head and	The majority of patients found APCD to be comfortable, effective, and well-tolerated, with one treatment showing statistically	The treatment was found to be safe, easy to use, and well-tolerated, showing edema reduction after just a single initial treatment	44	secondary	H&N	Single session	32m	N/A	once	N/A	N/A	N/A	NRCT, Prospective cohort study	Patient-reported comfort, feelings post-treatment, and likelihood of home use.		
Safety, Efficacy	Ridner 2010	Advanced pneumatic therapy in self-care of	Significant reduction in trunk circumference but the reduction in trunk circumference	100% compliance with IPC, Flexitouch used, study period was short	12	secondary	Trunk	10 days	1h		7 days per week	N/A	N/A	Flexitouch	Prospective cohort study	Patient reported symptoms, and trunk circumference		

D). SUPPORTIVE STUDIES ON COMPRESSION PRESSURE

Efficacy, best settings	Zaleska 2017	Indocyanine green near-infrared lymphangiography for evaluation of	Realtime movement of edema fluid observed using various compression modalities. Threshold pressures necessary	Determined sleeve pressure required for moving fluid proximal 40 mm Hg in the tissues which can be generated with was	N/A		UE, LE	single session						50 to 120 mm Hg gradient	N/A	Experimental study, single session	ICGL before and after, also measured tissue pressure using four different	
Other	Partsch 2011	Dose finding for an optimal compression pressure to reduce chronic edema of the extremities	The study examines the change in limb volume and circumference after 2 hours of using compression garments with different pressures, revealing that the upper limit for	Indirect estimate of required compression pressure	36 arm patients and 42 leg patients	secondary	UE, LE	N/A	N/A		N/A	N/A	N/A	N/A	N/A	Prospective cohort study	Limb volume and circumference before and after treatment	

ADL, activities of daily living; AE, adverse events; APCD, advanced pneumatic compression device; ASSES, American Shoulder and Elbow Surgeons; BCRL, breast cancer-related lymphedema; CB, compression bandages; CDT, complex decongestive therapy; CL, contralateral; FU, follow up; ggs, groups; h/hr, hour; H&N, head and neck; ICGL, indocyanine green; ICGL, indocyanine green near-infrared lymphography; IPC, intermittent pneumatic compression; its, joints; LE, lower extremity; LLCS, leg lymphedema complexity score; LYMQOL, Lymphedema Quality of Life scores; LSG, lymphoscintigraphy; LV, limb volume; LVA, lymphatic venous anastomosis; m, minutes; MLD, manual lymphatic drainage; MOA, mechanism of action; NIRFLI, near-infrared fluorescence lymphatic imaging; NP-PCD, non-programmable pneumatic compression device; NRCT, non-randomized control trial; PC, pneumatic compression; PCD, pneumatic compression device; PCT, pneumatic compression therapy; PMLE, post-mastectomy lymphedema; P-PCD, programmable pneumatic compression device; PREV, percentage reduction in edema volume; pts, participants; QOL, quality of life; RCT, randomized control trial; ROM, range of motion; s, seconds; IPC, intermittent pneumatic compression; SF-36, short form 36; SLD, self-lymphatic drainage; SPC, sequential pneumatic compression; TDC, tissue dielectric constant; TF, tissue fluid; UE, upper extremity; USG, ultrasound sonography test; w, weeks; yr, year.