

# A systematic review and meta-analysis of elastic stockings for prevention of thrombosis after orthopedic surgery

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**Background:** To investigate the preventive effect of elastic stockings on deep vein thrombosis (DVT) after orthopedic surgery by literature search and meta-analysis.

**Methods:** PubMed, Embase and Cochrane were selected as the search database platforms to search the literature of randomized controlled trials related to elastic stockings and DVT published from 2008 to date. Revman 5.3.5 software was used for statistical analysis of the data to obtain forest and funnel plots.

**Results:** In this study, 90 studies were initially screened and 7 were finally included, covering a total of 3,116 patients. Meta-analysis showed that the 7 studies had statistical heterogeneity ( $I^2=32\%$ , P=0.18), so a random effect model was used. The obtained statistic was [odds ratio (OR) =0.59, 95% confidence interval (CI): (0.34, 1.03)], the statistical effect size was Z=1.84, P=0.07, and the difference was not statistically significant, so a stepwise sensitivity analysis was performed by the exclusion method. One study was excluded, and the remaining 6 showed homogeneity ( $I^2=0\%$ , P=0.46). They were analyzed by subgroup according to the type of operation: ankle surgery or hip and knee arthroplasty. The internal literatures of each subgroup were homogeneous: ankle surgery subgroup ( $I^2=0\%$ , P=0.43), hip and knee arthroplasty subgroup ( $I^2=0\%$ , P=0.88). Therefore, fixed effect mode analysis was used, and the effect size of elastic stockings after ankle surgery was Z=3.65, P=0.0003, while the effect size of elastic stockings in the hip and knee arthroplasty subgroup was Z=1.23, P=0.22.

**Discussion:** Elastic stockings had an obvious preventive effect on DVT in patients undergoing ankle surgery, but not in patients undergoing lumbar, knee or spinal surgery. It is necessary to combine anticoagulant drugs and other physical therapies to prevent DVT.

Keywords: Deep venous thrombosis (DVT); elastic stockings; meta-analysis; orthopedics

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## Introduction

Deep venous thrombosis (DVT) occurs when the blood flow rate slows, or the blood becomes hypercoagulable, which makes the venous blood clot in the blood vessels and can stimulate post-thrombotic syndromes (PTS) such as tissue edema, ulcers, and varicose veins (1). Anesthesia of the orthopedic surgery can slow down the venous flow rate, formating a hypercoagulable state of blood, and after surgery the patients need limb immobilization for recovering, which is also an important cause of postoperative DVT (2,3). The incidence of DVT after orthopedic surgery has been reported to be between 1% and 25% (4). Therefore, prevention of postoperative thrombosis is very important, the use of anticoagulant drugs can prevent the formation of new thrombus and dredge the vein that has formed thrombus, in addition, some physical therapies can help to inhibit the growth of thrombus, such as leg elevation, intermittent compression inflation, plantar venous pump, and compression elastic stockings (5). Medical elastic stockings are used to contract the muscles with progressive pressure and squeeze venous blood back to the heart to prevent venous congestion and thus DVT (6). However, the results of such studies vary and the preventive effect of elastic stockings on thrombosis still remains unclear. Thus in the present study we included the latest studies in recent years for a meta-analysis to further clarify the preventive effect of elastic stockings. We present the following article in accordance with the PRISMA reporting checklist (available at https://dx.doi.org/10.21037/apm-21-2231).

## Methods

## Search strategy

PubMed, Embase and Cochrane were selected as the search platforms, using the keywords "stockings" AND "deep vein thrombosis" or "compression stockings" AND "DVT", or "Elastic stockings" AND "DVT" as the search criteria. The publication time and study type were limited to randomized control trials (RCTs) published from 2008 to date.

## Inclusion and exclusion criteria

## **Inclusion criteria**

(I) Study type: RCT of interventional nature, single- or multi-center study. (II) Study subjects were all patients undergoing orthopedic surgery (ankle surgery, hip arthroplasty, knee arthroplasty, spinal surgery, etc.). The patients were aged over 18 years, and did not have lower extremity DVT before surgery. (III) The study group must contain the elastic stockings and control groups. All patients took the same basic prevention method: anticoagulant drugs were given to high-risk patients with DVT, and asymptomatic patients did not need drug treatment, but ultrasound was given every 1 week. On this basis, the patients in the experimental group were given elastic socks physical therapy, whereas the control group was limited to a placebo medical cylindrical bandage or no preventive measures. (IV) The study clearly described the DVT diagnostic method (biphasic ultrasound), diagnostic criteria and DVT index data.

#### **Exclusion criteria**

(I) Non-randomized, non-concurrent controlled study, observational case-control study, cross-sectional study, review, guideline, investigation, case analysis, systematic review. (II) The study did not specify the orthopedic surgery, or the study subjects were mixed with the study including gynecological, cardiac surgery and other surgical patients. (III) The study subjects had been diagnosed as DVT patients before surgery. (IV) The study used elastic stockings as a treatment but not as a preventive measure.

## Risk of bias assessment

We evaluated the risk of bias in the studies from six aspects according to the criteria in the Cochrane Handbook for Systematic Reviews of Interventions (7): the generation of random sequences, classification concealment, blind method, whether outcome assessment was incomplete, selective reporting, and other biases.

#### Literature screening

Two researchers searched the databases, combined the search results, read the abstracts of each study as the primary screening, read the full text of the study after primary screening, continued screening according to the inclusion and exclusion criteria, and evaluated the quality of the studies for inclusion in the analysis, as shown in *Figure 1*.

## Data collection

The researchers read the full text of each of the included articles, and extracted the following information: authors, year of publication, the number of interventions, sex ratio,

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Consider, if feasible to do so, reporting the number of records identified from each database or register searched (rather than the total number across all databases/registers).

If automation tools were used, indicate how many records were excluded by a human and how many were excluded by automation tools.

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Figure 1 Flow chart of literature search and screening. RCT, randomized control trial.

age, surgical procedure, intervention methods for study and control groups, and the incidence of DVT. The data were organized into tables for further analysis.

#### Statistical analysis

We used Revman 5.3.5 software released by the Cochrane Collaboration for comprehensive analysis, the statistical method was Mantel-Haenszel (M-H) and the odds ratio (OR) and 95% confidence interval (CI) were used to report the incidence of DVT in the two groups of intervention methods, with P<0.05 considered statistically significant. The software created forest plots to display the results. All studies with DVT incidence were included in the study and the index expressed by % in all studies was converted into

actual numbers of cases. The heterogeneity of the studies was analyzed by  $I^2$  analysis and Q validation, and  $I^2>25\%$  or P<0.1 indicated heterogeneity of results. If heterogeneity among the studies was suggested, sensitivity analysis was performed using literature analysis and a case-by-case elimination method, and a funnel plot was used to represent publication bias.

#### **Results**

#### Literature screening results

We initially screened 90 studies and based on our inclusion and exclusion criteria and literature quality evaluation, 7 were finally included, covering a total of 3,116 patients (*Table 1*).

Authors	Year	Number (T/C)	Male (T/C)	Mean age (years) (T/C)	Surgery	Intervention group mode	Control group mode	DVT events (T/C)
Sultan <i>et al.</i> (8)	2014	44/46	25/29	46.4/47.0	Ankle surgery	Compression stockings	TubiGrip Medical Cannulated Bandage	5/10
Winge <i>et al.</i> (9)	2018	82/71	28/29	57.0/61.0	Ankle surgery	Compression stockings	TubiGrip Medical Cannulated Bandage	14/31
Chin <i>et al.</i> (10)	2009	110/110	14/9	65.0/67.0	Total knee arthroplasty	Graded compression stockings	No precautions	14/22
Kim <i>et al.</i> (11)	2015	40/50	NA	NA	Hip/knee arthroplasty	Graded compression stockings	No precautions	2/3
Maki <i>et al.</i> (12)	2021	595/164	234/76	56.0/58.0	Hip/knee arthroplasty	Compression stockings	No precautions	2/0
Majeed <i>et al.</i> (13)	2013	1,040/984	NA	NA	Hip/knee arthroplasty	Compression stockings	No precautions	5/5
Takahashi <i>et al.</i> (14)	2012	1,434/541	NA	NA	Spinal operation	Compression stockings	No precautions	8/3

Table 1 Baseline characteristics of the included studies of DVT

DVT, deep vein thrombosis; T, intervention group; C, control group; NA, not available.

Table 2 Bias assessment b	based on	Cochrane	systematic	review	criteria	for rand	lomized	intervo	entions
			~						

Study	Generation of random sequence	Classification hiding	Blind method	Inadequate outcome assessment	Optional reporting	Other bias
Sultan <i>et al.</i> (8)	Yes	Yes	Yes	Yes	Unclear	Unclear
Winge et al. (9)	Yes	Yes	Yes	Yes	Yes	Unclear
Chin <i>et al.</i> (10)	Yes	Yes	Unclear	Yes	Yes	Unclear
Kim <i>et al.</i> (11)	Unclear	Yes	Yes	Yes	Unclear	Unclear
Maki e <i>t al.</i> (12)	Yes	Yes	Yes	Yes	Yes	Unclear
Majeed et al. (13)	Yes	Unclear	Yes	Yes	Yes	Unclear
Takahashi et al. (14)	Yes	Yes	Yes	Yes	Yes	Yes

## Risk of bias assessment

The bias of the 7 studies was assessed according to Cochrane Randomized Intervention Evaluation System (*Table 2*).

## Meta-analysis results

## Statistical analysis

Meta-analysis showed that the 7 studies had statistical heterogeneity ( $I^2=32\%$ , P=0.18), so a random effect model was used, and the obtained statistics were [OR =0.59, 95% CI: (0.34, 1.03)]. The statistical effect value was Z=1.84,

P=0.07, and the difference was not statistically significant, as shown in *Figure 2*. The risk of literature bias may bias the results.

### Sensitivity analysis

Due to heterogeneity in the 7 articles, the study by Takahashi *et al.* (14), which described spinal surgery, was excluded by the case-by-case exclusion method, and the remaining 6 studies showed homogeneity ( $I^2=0\%$ , P=0.46). These studies were analyzed by subgroup according to the type of operation: ankle surgery or hip and knee arthroplasty. The internal validity of each subgroup showed homogeneous: ankle surgery subgroup ( $I^2=0\%$ , P=0.43),

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	Compression sto	ckings	NO Compression stoc	kings		Odds Ratio		Odds	Ratio	
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl		M-H, Rande	om, 95% Cl	
Chin P 2009	14	110	22	110	26.4%	0.58 [0.28, 1.21]			-	
Kim 2015	2	40	3	50	7.8%	0.82 [0.13, 5.19]				
Majeed H 2013	5	1040	5	984	14.3%	0.95 [0.27, 3.28]				
Maki S 2021	2	595	0	164	3.2%	1.39 [0.07, 29.01]			•	
Sultan MJ 2014	5	44	10	46	15.7%	0.46 [0.14, 1.48]			-	
Takahashi H 2012	11	1434	1	541	6.5%	4.17 [0.54, 32.41]			•	
Winge R 2018	14	82	31	71	26.0%	0.27 [0.13, 0.56]				
Total (95% CI)		3345		1966	100.0%	0.59 [0.34, 1.03]		•		
Total events	53		72							
Heterogeneity: Tau <sup>2</sup> =	0.17; Chi <sup>2</sup> = 8.88, 0	0.18); I <sup>2</sup> = 32%				-	1			
Test for overall effect:	Z = 1.84 (P = 0.07)						0.01	U.1	1 1U	100
								Favours [GC]	Favours [NO GC]	

Figure 2 Forest plot of use of elastic stockings for the prevention of DVT after orthopedic surgery. CI, confidence interval; DVT, deep vein thrombosis.

	Compression stockings		NO Compression stockings		Odds Ratio	Odds Ratio			
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% CI		
1.1.1 ankle									
Sultan MJ 2014	5	44	10	46	13.6%	0.46 [0.14, 1.48]			
Winge R 2018	14	82	31	71	43.2%	0.27 [0.13, 0.56]			
Subtotal (95% CI)		126		117	56.7%	0.31 [0.17, 0.58]	◆		
Total events	19		41						
Heterogeneity: Chi <sup>2</sup> =	0.61, df = 1 (P = 0.4)	3); I <sup>2</sup> = 09	6						
Test for overall effect:	Z = 3.65 (P = 0.0003	3)							
1.1.2 hip/knee									
Chin P 2009	14	110	22	110	30.1%	0.58 [0.28, 1.21]			
Kim 2015	2	40	3	50	4.0%	0.82 [0.13, 5.19]			
Majeed H 2013	5	1040	5	984	8.0%	0.95 [0.27, 3.28]			
Maki S 2021	2	595	0	164	1.2%	1.39 [0.07, 29.01]			
Subtotal (95% CI)		1785		1308	43.3%	0.70 [0.39, 1.24]			
Total events	23		30						
Heterogeneity: Chi <sup>2</sup> =	0.69, df = 3 (P = 0.8)	8); 17 = 09	6						
Test for overall effect:	Z = 1.23 (P = 0.22)								
Total (95% CI)		1911		1425	100.0%	0.48 [0.31, 0.73]	•		
Total events	42		71						
Heterogeneity: Chi <sup>2</sup> = 4.66, df = 5 (P = 0.46); i <sup>2</sup> = 0%									
Test for overall effect; Z = 3,44 (P = 0,0006)									
Test for subgroup differences: Chi <sup>2</sup> = 3.38. df = 1 (P = 0.07). I <sup>2</sup> = 70.4%									

Figure 3 Forest plot of subgroups of elastic stocking use for the prevention of DVT after orthopedic surgery. CI, confidence interval; DVT, deep vein thrombosis.



**Figure 4** Funnel plot of use of elastic stockings for prevention of DVT after orthopedic surgery. DVT, deep vein thrombosis.

hip and knee arthroplasty subgroup ( $I^2=0\%$ , P=0.88). Using fixed effect mode analysis, the effect size of elastic stockings after ankle surgery was Z=3.65, P=0.0003, suggesting that the use of elastic stockings can prevent the occurrence of

DVT, whereas the effect size of elastic stockings in the hip and knee arthroplasty subgroup was Z=1.23, P=0.22, suggesting no significant effect in preventing the occurrence of DVT (*Figure 3*).

#### Analysis of publication bias

The funnel plot showed that the two subgroups were unevenly distributed, suggesting the presence of publication bias (*Figure 4*).

## Discussion

Lower extremity DVT is an abnormal blood coagulation state that can cause vascular intimal lesions, causing varying degrees of PTS, and even the risk of pulmonary embolism (PE) (15). Medical elastic stockings are a physical therapy measure that produce a stepped decrease in pressure from the ankle, which reduces blood viscosity, promotes blood return, increases flow rate, reduces blood stasis, and protects the vascular endothelium, thereby achieving the effect of preventing DVT (16). However, its application effect is still under discussion; some studies (8,10-12) have shown an obvious preventive effect on DVT, while some studies (9,13,14) believe the preventive effect is not significantly different from that of conventional measures. In this study, a total of 3,116 orthopedic surgery patients in 7 articles were screened for a meta-analysis, and the study results showed that in the random effect mode, the OR of DVT was [0.59, 95% CI: (0.34, 1.03)] when elastic stockings were used versus no elastic stockings, but the statistic value was Z=1.84, P=0.09, suggesting that the difference was not statistically significant; that is, elastic stockings had no significant preventive effect on DVT. This is different from meta-analyses performed by Sachdeva et al. (17) and Hajibandeh et al. (18), and may be due to the fact that we focused on patients after orthopedic surgery, whereas the subjects of the other meta-analysis were patients with different surgeries. Our result suggests that for patients undergoing orthopedic surgery, it is not enough to use elastic stockings alone for DVT prevention, and it needs to be performed in combination with other methods such as anticoagulant drugs, pneumatic compression, etc., in order to achieve better results.

In this study, the 7 articles showed heterogeneity  $(I^2=32\%, P=0.18)$ , so the case-by-case exclusion method was used for the sensitivity analysis. The results showed that after removing the study (14) whose subjects were patients undergoing spinal surgery, the remaining 6 showed homogeneity ( $I^2=0\%$ , P=0.46), so they were analyzed according to the surgical procedure. The results showed homogeneity among the internal literatures, while the heterogeneity between the subgroups was higher ( $I^2=70.4\%$ ), suggesting that the benefit of using elastic stockings in patients undergoing ankle surgery was different from that in patients undergoing hip and knee surgery. The statistical value showed that the effect size of using elastic stockings after ankle surgery was Z=3.65, P=0.0003, suggesting that the use of elastic stockings can effectively prevent the occurrence of DVT; while the effect size of using elastic stockings after hip and knee arthroplasty was Z=1.23, P=0.22, suggesting no significant effect on preventing the occurrence of DVT. A study by Ayhan et al. (19) showed that there was a difference in patient comfort between elastic stockings for the knee and those for the hip, which may affect patient compliance. The types of elastic stockings used after different orthopedic surgeries differ, so their preventive effect on DVT may also differ. In addition, for

the surgeries with a wide range of effects, the possibility of postoperative DVT in patients is higher, and the preventive effect of elastic stockings is limited, so should be combined with other anticoagulant methods.

In this study, the publication bias analysis was performed for the 6 studies remaining after excluding the study by Takahashi et al. (14). Our analysis found significant asymmetry on both sides of the funnel plot, suggesting there was a risk of publication bias. One possible reason was that people overestimated the effect of elastic stockings. In addition, there are various types of elastic stockings that produce different pressures. One study (20) showed that graded elastic stockings had the best preventive effect when the pressure was 18 mmHg at the ankle, 14 mmHg at the lower leg and 8 mmHg at the knee. We did not distinguish the specific types of elastic stockings used in the different studies. In addition, we did not record the intervention duration of elastic stockings in the studies, which varied from 2 weeks to 3 months. In a study conducted by Health Quality Ontario (21), wearing a compress stocking may be an effective way of preventing chronic venous and it's costeffective for the long run, and it could help to reduce the recurrence of DVT by half.

All the studies did not report side effects of compress stockings, but as Robertson BF (22) reported, for those who are at greater risk of skin damage, the application of compress stockings need careful assessment and close monitoring.

One study (23) showed that for patients with moderate or high risk of DVT before elective surgery, pharmacological thromboprophylaxis alone was not inferior to the combination of pharmacological thromboprophylaxis and compression stockings, which means anticoagulant drugs still should be the dominant way to preventing DVT.

#### Conclusions

In this meta-analysis of the preventive effect of elastic stockings on DVT after orthopedic surgery, a total of 3,116 patients in 7 studies were included. The results showed that elastic stockings had a significant preventive effect on DVT in patients undergoing ankle surgery, but not in patients undergoing lumbar, knee or spinal surgery. It is necessary to combine anticoagulant drugs and other physical therapy methods to jointly prevent DVT.

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## Footnote

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