Anatomical Consideration of Deep Calf Veins: Application to Catheter-Directed Thrombolysis

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Research Article

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Abstract

An antegrade approach is frequently used in catheter-directed thrombolysis to remove deep-vein thrombosis. However, the antegrade approach is difficult when accessing veins with small diameters, therefore, understanding the variation of deep calf vein is important. This study measured the diameters and surface areas of the proximal and distal posterior tibial vein, peroneal vein, and anterior tibial vein in order to determine which are preferable for venous access. This study dissected 132 legs from Korean and Thai cadavers. The proximal and distal posterior tibial vein, peroneal vein, and anterior tibial vein were scanned and measured. The mean diameter and surface area were largest for the proximal tibial vein, at 6.34 mm and 0.312 cm$^2$, respectively, followed by the anterior tibial vein (5.22 mm and 0.213 cm$^2$), distal posterior tibial vein (3.29 mm and 0.091 cm$^2$), and peroneal vein (3.43 mm and 0.081 cm$^2$). The proximal posterior tibial vein and anterior tibial vein have large diameters and surface areas, which make them ideal for applying an antegrade approach in catheter-directed thrombolysis. The distal posterior tibial vein and peroneal vein are not recommended due to their smaller surface areas and also the anatomical variations therein.

Introduction

Deep-vein thrombosis (DVT) is a frequent vascular disorder that is associated with high morbidity. The prevalence of DVT has increased to 0.1% in the general population, which is largely attributable to advances in diagnostic methods such as ultrasound scanning$^1$. DVTs occur most frequently in the lower legs$^2$, and 20% of untreated DVTs advance to proximal veins and cause a pulmonary embolism$^3,4$. Many clinicians suggest practitioners should consider anatomical variations as a risk factor for DVT. Thereby, previous study identified and classified the anatomical variation in the deep calf veins (Figure 1)$^5$.

Administering a systemic thrombolytic agent is the conventional therapy for reducing the risk of pulmonary embolism and DVT reoccurrence. However, applying only a systemic thrombolytic agent is associated with post-thrombotic syndrome (PTS) in 20–50% of treated patients$^6$. PTS should be avoided since it is a chronic problem of DVT that includes ulceration, swelling, and dermatosclerosis$^7$.

Systemic thrombolytic agents should also not be applied alone to patients with high blood pressure, stroke, and arterial aneurysms$^6$. In the last few decades, complementary and combination therapies for catheter-directed thrombolysis (CDT) have been suggested that involve the selective release of thrombolytic agents using intervention catheters. Previous studies have demonstrated that CDT is an effective and safe treatment option for DVT$^8,9$. The combination therapy of CDT and systemic thrombolytic agents for thrombus elimination is recommended in patients with acute DVT in order to relieve acute symptoms and prevent PTS$^{10}$. The American Heart Association recommended CDT as the primary treatment for patients with acute DVT and for preventing symptom progression and the rapid extension of a thrombus, followed by anticoagulation therapy$^{11}$. 
CDT has the benefits of the rapid relief of painful symptoms with a smaller amount of thrombolytic agent\textsuperscript{12,13}. The catheter is inserted proximal or distal to the affected sites in the retrograde or antegrade venous approach, respectively\textsuperscript{14}. The venous entrance points of CDT are the posterior tibial vein, anterior tibial vein, popliteal vein, and femoral vein. The venous entrance point is selected by considering anatomical variations in and the diameter of the accessing vein. It is necessary to carefully select the venous entrance point by taking into consideration all of the anatomical pros and cons.

A retrograde venous approach via the femoral vein or popliteal vein has the advantages of a short procedural time due to the large diameter of the vein and the smaller number of anatomical variants. The disadvantages of this approach are the possibility of valvular injury due to the access direction being against the direction of blood flow, insufficient removal of the thrombus, and difficulty of positioning the patient during the procedure\textsuperscript{15}.

An antegrade venous approach via the anterior tibial vein, posterior tibial vein, or peroneal vein has the advantages of sufficient thrombus clearing, easier positioning, and less valvular damage. Additionally, the antegrade approach has no limitation in clearing the proximal extent of the thrombus, including in cases of an inferior popliteal thrombus, which are often not resolved\textsuperscript{15}. The disadvantages of the antegrade approach are the long procedural time needed to obtain access and evaluate veins with small diameters and variations\textsuperscript{15}.

Since CDT is a fairly new treatment, physicians must focus on more-effective and safer access points and approaches. The diameters of these veins and variations therein should be carefully considered, since CDT is associated with bleeding complications at the insertion point. The venous anatomy of the distal extremities is significantly more unpredictable and complex than those of the corresponding arteries and veins in proximal regions\textsuperscript{16}.

A previous study revealed the diameters of the popliteal veins that are commonly used for the retrograde approach of CDT\textsuperscript{17}. However, no previous study has investigated the diameters of deep calf veins.

This study identified the diameters and surface areas of the posterior tibial vein, anterior tibial vein, and peroneal tibial vein. Knowledge of the diameters and surface areas of these veins will help practitioners to select the best vein when adopting an antegrade approach in CDT.

**Methods**

This study was conducted in accordance with the principles outlined in the Declaration of Helsinki. Informed consent and approval were obtained from the families of the cadavers before the dissections were performed. All cadavers used in this study were legally donated and approved from ethics committee of the Surgical Anatomy Education Center, Yonsei University College of Medicine (approval code 19-003; approval date: May 15th, 2019).
This study dissected 132 legs from Korean and Thai cadavers, comprising 42 males and 24 females with a mean age of 69.3 years and an age range 43–96 years. The proximal and distal posterior tibial vein, peroneal vein, and anterior tibial vein were sliced at 2 cm from their diverging or converging points (Figure 3). The diameter was determined as the mean values of the measured longitudinal and horizontal values.

Proximal and distal posterior tibial veins were defined in accordance with previous research \(^5\). According to Yi & Kim, the proximal tibial vein begins from the point where the distal posterior tibial vein and peroneal vein merge, and it turns into the popliteal vein at the point where the anterior and proximal tibial veins merge (Figure 4). The distal tibial vein starts from the point where the medial and lateral plantar veins merge, and it turns into the proximal posterior tibial vein at the point where it merges with the peroneal vein (Figure 2).

Sliced specimens were mounted and then scanned (CanoScan LiDE 220), and their diameters and surface areas were measured using standard imaging software (Image J, NIH, Bethesda, MD) (Figure 5).

**Statistical analysis**

Comparisons among the cadavers based on sex were performed using independent t tests, with p values of <0.05 considered to indicate significant differences. All measured values were analyzed using SPSS software (version 15.0).

**Results**

**Diameters**

The popliteal, proximal posterior tibial, anterior tibial, peroneal, and distal posterior tibial veins had diameters of 7.42±1.21 mm (mean±SD, range 5.43–9.58 mm), 6.34±1.51 mm (range 3.3–8.7 mm), 5.22±1.38 mm (range 3.40–7.43 mm), 3.43±1.24 mm (range 1.53–5.64 mm), and 3.29±1.09 mm (range 1.23–5.37 mm), respectively (Figure 2). There were no sex-related differences in the diameters.

**Surface area**

The popliteal vein had the largest surface area, at 0.432±0.092 cm\(^2\), followed by the proximal posterior tibial vein (0.312±0.093 cm\(^2\)), anterior tibial vein (0.213±0.091 cm\(^2\)), peroneal vein (0.091±0.062 cm\(^2\)), and distal posterior tibial vein (0.081±0.065 cm\(^2\)).

**Discussion**

The main objective of DVT treatment is to inhibit the expansion of the thrombus load within the affected vein and avoid subsequent pulmonary embolism. PTS also needs to be prevented, which is triggered by valvular insufficiency and an inadequately treated or untreated venous obstruction \(^18,19\).
Previous research has shown that a residual thrombus and recurrent DVT are powerful predictors of subsequent PTS. It was demonstrated that there is a direct correlation between insufficient thrombus lysis and the long-term development of PTS. Another study demonstrated that improving the venous valvular function by resolving thrombus can reduce the probability of PTS.

Conventional treatment with systemic anticoagulation was found to result in problems with residual thrombus, and so the treatment standard was changed to combination therapy with CDT. Combined treatment with systemic anticoagulation and CDT resulted in fewer remaining thrombi and reduced the recurrence of thrombosis, while at the same time decreasing the prevalence of PTS. Fleck et al. proposed that combined therapy with CDT is more advantageous in younger patients and those with a more substantial thrombus load.

The popliteal vein is currently the most frequently utilized for CDT. However, the retrograde approach of CDT via the popliteal vein has a reported downside of residual thrombus after CDT, particularly in the inferior popliteal vein. Furthermore, the patient needs to be in the prone position during the retrograde approach, and the associated positional difficulties are particularly problematic in the presence of severe obesity, paralysis, or fractures.

This situation has led to the antegrade approach being commonly used as an alternative method from the posterior tibial vein. A previous clinical study demonstrated that there were more sequelae during short-term follow-up for enhanced lysis of the distal segments of a thrombosis when the approach was from the posterior tibial vein. Since the approach is in the direction of the blood flow and results in less damage to the valves and vein walls, the posterior tibial vein was suggested as an best venous entry point for CDT.

Bendix et al. reported that the long-term outcomes were better for posterior tibial venous access than for popliteal venous access. However, those authors indicated that great care is required when performing antegrade approaches due to the smaller diameter of the veins. Our research has revealed that the small diameter of the distal posterior tibial vein limits its utilization, and so we recommend the proximal posterior tibial vein as the best venous entry point.

Despite antegrade approaches having many advantages over retrograde approaches, they have impediments for clinicians associated with variations in the posterior tibial vein. Yi & Kim recently classified the posterior tibial vein into the proximal posterior tibial vein and the distal posterior tibial vein based on significant anatomical differences regarding the merging of other veins. The distal posterior tibial vein merges with the peroneal vein and turns into the proximal posterior tibial vein, while the proximal posterior tibial vein merges with the anterior tibial vein and becomes the popliteal vein. Most veins have two accompanying veins in addition to the corresponding artery. However, those authors found that the distal posterior tibial vein had two or three accompanying veins, the proximal posterior tibial veins had one or two accompanying veins, the anterior tibial vein had two accompanying veins, and the peroneal vein had two accompanying veins.
In conclusion, CDT should be carried out by trained physicians with sound knowledge of the venous structures as well as the diameters of the approached veins. This study measured the surface areas of deep calf veins in order to identify the most-appropriate veins for CDT. The obtained results suggest that the distal posterior tibial vein is not appropriate for CDT due its larger anatomical variations and smaller surface area compared with other veins. We also suggest that the peroneal vein should not be used, since it has the second-smallest surface area of all the deep calf veins. We further suggest that the proximal tibial vein has fewer variations and that the anterior tibial vein does not have the variations reported previously.

The present findings indicate that a thorough understanding of the variations and diameters of deep calf veins is necessary for guiding practitioners who are selecting an antegrade approach for CDT.

**Abbreviations**

Deep-vein thrombosis (DVT), post-thrombotic syndrome (PTS), catheter-directed thrombolysis (CDT)

**Declarations**

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**Author Contribution**

KHY: research concept, study design, visualization and writing of the manuscript

HJK: reviewing/editing a draft of the manuscript, supervise the manuscript

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Conflicts of interest/Competing interests

The author(s) declare no competing interests.

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**Figures**

**Figure 2**

Image of the left side of a deep calf vein and an artery specimen.
Ultrasonographic image of the deep calf veins, showing the PPTV (A) merging with the ATV (B) to form the PoV. The popliteal artery (PoA) (C) is present deep to the veins. P, posterior; A, anterior; M, medial; L, lateral.