Varicose vein ablation: navigating the treatment options

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Editorial

Nick Morrison, MD

Over the past two decades, technological advances in the diagnosis and treatment of chronic superficial venous disorders have resulted in traditional, more aggressive surgical methods falling out of favor in many parts of the world. Thankfully, gone are the days of massive leg bandages and prolonged bed rest following surgery. Yet, improvements in those surgical techniques, especially groin-to-knee invagination stripping under ultrasound-guided local tumescent anesthesia and ambulatory phlebectomy through microincisions, have significantly improved the patient's treatment experience.

Today, surgical patients can expect a far more comfortable recovery and an earlier return to normal activities. These surgical techniques must not be dismissed as archaic, because, in the hands of a meticulous surgeon, they provide a very cost-effective treatment, with anatomic and patient-reported outcomes on par with newer endovenous techniques.

That said, dramatic improvements in duplex ultrasound have led to enhanced understanding of the pathophysiology of venous disease, a development that has allowed for less invasive treatment of superficial venous incompetence. Although lacking enthusiastic adoption by most venous surgeons, saphenous-sparing principles and techniques are intriguing and valuable components in the armamentarium of thoughtful practitioners.

With these advances have come challenges to dogmatic, traditional beliefs about venous disease diagnosis and treatment. Evidence-based principles and guidelines have emerged to become as important as or even more important to practitioners than their own clinical experience.

Michel Perrin's treatise on endovenous therapeutic modalities presents a timely overview of these emerging and increasingly popular methods. Evaluation of randomized controlled trials of endothermal ablation techniques can be problematic because some studies may have used first-generation radiofrequency technology and shorter wavelength lasers, both of which will skew results. Longer wavelength laser generators and technological advances in laser fiber and radiofrequency catheters have largely overcome disadvantages of older radiofrequency and laser technology, so one has to evaluate endothermal randomized controlled trials with these concepts in mind.

Furthermore, analysis of cost effectiveness of different procedures may include only the medical costs of the actual procedure rather than taking into account cost savings to society as a result of a more rapid return to work and normal activities.

So-called nonthermal, nontumescent treatments are emerging techniques that will further minimize the impact of treatment on patients' daily life. Foam sclerotherapy has been recognized as a very cost-effective modality. And even though anatomic results may not appear to be as good as with endothermal and surgical treatment, foam sclerotherapy is easily repeatable in cases of incomplete success. However, because of the use of different sclerosing agents and gases and the myriad foam production and delivery protocols, comparison of outcome data from studies using foam sclerotherapy are nearly impossible. Caution is advised when attempting to draw definitive conclusions from foam sclerotherapy studies. Earlier concerns about severe, life-altering neurosensory events following foam sclerotherapy have subsided as the true incidence has been shown to be extremely rare. To the contrary, very serious adverse events are regularly reported with the use of illegal or nonapproved sclerosants, resulting in sometimes severe consequences for the patient and medico-legal exposure for the practitioner.

The use of posttreatment compression therapy has been the subject of controversy with the less invasive treatment methods. Never very strong, evidence for the use of compression stockings in the management of venous disease has recently been challenged, with further studies to clarify the role of compression stockings in development or currently being conducted.

Finally, recurrence is a constant bedevilment for nearly all patients with venous disorders. Treatment modalities have surely improved the patient experience. However, with our current understanding of the pathophysiology of superficial venous disorders, recurrence is considered nearly inevitable with the incidence probably about the same as recurrence following traditional surgical intervention, albeit by different mechanisms. Therefore, before we pat ourselves on the back for our accomplishments, perhaps some perspective is in order.

Plus ça change, plus c'est la même chose – Jean-Baptiste Karr

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An update on operative treatments of primary superficial vein incompetence

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Abstract

For more than a century, open surgery and liquid sclerotherapy were the only options used for operatively treating primary varices. In the last 20 years, management of primary varices has dramatically changed due to ultrasound investigations and innovative techniques. Development of endovenous treatments, including thermal ablation and/or chemical ablation, has provided a patient-friendly option for an office-based procedure, improving both the postoperative course and convalescence duration. This book will describe the new procedures and their possible complications, provide an analysis of the outcomes after the new procedures for short-, mid-, and long-term follow-up, and discuss the indications and the international guideline recommendations for operative treatment of varicose veins. For the outcome analysis, all randomized controlled trials (RCTs) published since 1990 on operative treatments of varicose veins were collected and the references were gathered in tables according to either the procedure used or the patient's clinical status. Case series and meta-analyses were taken into account in this review when RCTs were not available. For more details regarding clinical or instrumental outcomes of the studies described, please go to www.phlebolymphology.org.

Preface

The term operative treatment has been intentionally chosen instead of interventional treatment because interventional treatment means any kind of treatment that interferes with the natural history of the disease. For example, both compressive treatments and venoactive drugs modify the natural evolution of primary varicose veins.

Introduction

For a century, ancillary open surgery had the highest recommendation, and subsequently, was the most frequently used procedure for operatively treating varicose veins. In the past decade, the development of minimally invasive endovenous techniques for primary superficial venous reflux has provided a patient-friendly means of treating this disorder as an office-based procedure with ablation of the saphenous veins and tributary varicosities by using radiofrequency ablation, endovenous laser ablation, or sclerotherapy. Sclerotherapy regained favor for two reasons: (i) ultrasound investigation, which provided security for the

procedure; and (ii) the use of foam, which enhances the efficacy of the sclerosing agent. More recently, new procedures have been used, including steam ablation, ClariVein, laser-assisted foam sclerotherapy, and glue, and these procedures will be described in the present book.

Simultaneously, surgery, including the CHIVA procedure (Cure Hemodynamique de l'Insuffisance Veineuse en Ambulatoire [conservative ambulatory hemodynamic management of varicose veins]),[1] and more recently, the ASVAL procedure (ablation selective des varices sous anesthésie locale [ambulatory selective vein ablation under local anesthesia]),[2] were developed to preserve the great saphenous vein.

Open surgery without conservation of saphenous trunks

Modern open surgery should be performed under local anesthesia and directed by preoperative ultrasound assessment and skin mapping. Treatment of the great saphenous vein involves flush ligation of the saphenofemoral junction, which is completed using saphenous invagination stripping. Stripping can also be done using a cryoprobe. Treatment of the incompetent small saphenous vein usually involves flush saphenopopliteal junction ligation and stripping by invagination. Nontruncal varicosities can be excised using stab avulsion–powered phlebectomy or they can be treated with sclerotherapy in the same session or later.

Stripping of both the great saphenous vein below the knee and the distal small saphenous vein may reduce varicose vein recurrence, but it is associated with an increased risk of nerve injury.[3] The usefulness of flush ligation was recently called into question after a randomized controlled trial.[4] Nontruncal varicosities can be excised either by stab avulsion–powered phlebectomy, or treated by sclerotherapy in the same session or later.

In addition, there is a consensus for recommending elastic compression stockings for no more than 1 week after the operation.[5,6]

Complications of surgery

The early complications of surgery include discomfort (common), bruising (common), hematoma (rare), bleeding (very rare), lymphatic damage (rare), femoral vein or artery injury (extremely rare),[7] wound infections (2% to 6%), and injury of the saphenous or sural nerve (10%). Symptomatic and asymptomatic deep venous thrombosis and pulmonary embolism following open surgery vary from 0.4% to 5.3% and 0% to 0.5%, respectively.[8,9] The risk of complications, such as venous thromboembolisms, increase with redo surgery and surgery of the small saphenous vein.[8] Modern open surgery under local anesthesia has dramatically lowered the rate of thromboembolic complications. Late complications include permanent nerve damage (5%).[10]

Open surgery with preservation of the saphenous trunk

CHIVA

Due to the possible future use of the great saphenous vein as a vascular graft, it is necessary to preserve the vein.[1] The principle of the CHIVA technique consists of redistributing refluxes from the superficial to the deep system using staged ligations on the great saphenous vein or tributaries. CHIVA is a complex procedure that requires careful mapping and understanding of the anatomy and function of the superficial system by well-trained and experienced physicians who are aware of the shunt classifications.[11]

ASVAL

While CHIVA is based on a descending theory, the ASVAL method is based on an ascending or multifocal approach to the primary varicose veins. In order to improve or suppress the saphenous vein reflux, a stab phlebectomy of incompetent tributaries is performed to remove the distal venous reservoir. Compared with trunk varicose vein ablation, the major advantage of ASVAL is the preservation of the great saphenous vein. After the ASVAL procedure, most patients had less advanced stages of varicose veins.[2]

Endovenous thermal ablation

The term "endovenous thermal ablation" includes radiofrequency ablation, endovenous laser ablation, endovenous steam ablation, and endovenous microwave ablation. In endovenous thermal ablation procedures, ablation of the treated vein is achieved using heat, which is delivered into the vein through a percutaneously placed catheter or probe. The heat causes a direct thermal injury to the vein wall, resulting in destruction of the endothelium, denaturation of collagen in the media, and subsequently, thrombotic and fibrotic occlusion of the vein. Endovenous thermal ablation is performed under local tumescent anesthesia (except for endovenous microwave ablation) to provide anesthesia; protect the perivenous tissue from the heat created by the catheter, probe, or wire when activated; and spasm the vein to obtain the best contact with the heating device. In addition, all endovenous thermal ablation procedures are performed using ultrasound guidance and conducted as an outpatient-based procedure.

For the great saphenous vein, echo-guided vascular access occurs just below the knee (except for endovenous microwave ablation); therefore, heating is done from the groin (2 cm below the saphenofemoral junction) down to the distal part of the vein, usually just below or above the knee. For the small saphenous vein, echo-guided access occurs at the lower one-third of the lower leg, and heating is done from the popliteal fossa (2 cm below the saphenopopliteal junction) down to just above (8 to 10 cm) the tibial malleolus.

Radiofrequency ablation

Introduced in 2007, the current ClosureFAST radiofrequency catheter (VNUS Medical Technologies/Covidien) *(Figures 1 and 2)* is easy to use. The entire pullback time takes 3 to 4 minutes, generating heat around 120°C. Celon RFITT, another radiofrequency ablation system for bipolar radiofrequency-induced closure, is now available (Olympus Medical Systems). This system generates heat at 60 to 85°C and operates with a continuous pullback speed of 1 to 1.4 cm/second.
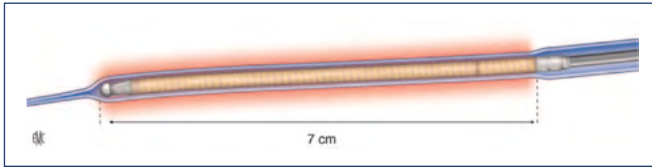
Figure 1. ClosureFAST catheter.
The first 7 cm (left) of the coated heating element and the thermocouple (right).
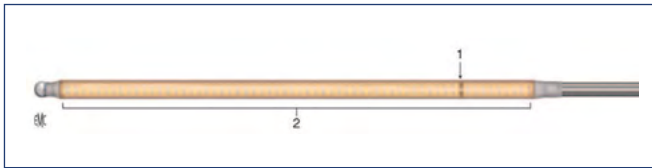
Figure 2. ClosureFAST heater and thermocouple catheter.
The thermocouple (1) is mounted distally to the heating element (2).

Endovenous laser ablation

Fiber lasers can provide either low wavelength beams (810, 940, and 980 nm) or high wavelength beams (1319, 1320, 1470, and 1500 nm). Theoretically, light of lower wavelengths is less specifically absorbed by the chromophores (hemoglobin, water, proteins) compared with the light of higher wavelength lasers.[12] Previously, the fibers were bare tipped, but the new radial fibers are more effective and include the Radial fiber R (Biolitec) *(Figure 3)*, Never-Touch R (Angiodynamics), and Tulip fiber R (Tobric). A continuous withdrawal technique is the current rule and it is recommended to deliver 50 to 70 J/cm of energy.
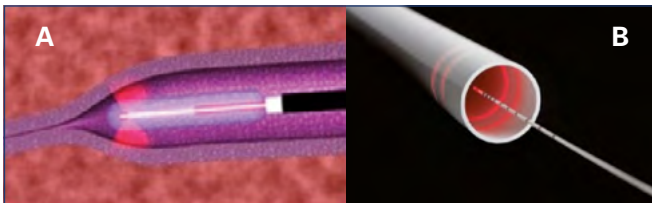
Figure 3. Fiber with radial emission.
Fiber with single radial emission (Panel A) and double radial emission (Panel B).

Radiofrequency ablation vs endovenous laser ablation

Endovenous laser ablation and radiofrequency ablation are similar techniques that treat similar patient profiles. After percutaneous access, the radiofrequency ablation catheter or laser fiber is pushed proximally until the tip is positioned 2 cm from the saphenofemoral junction or saphenopopliteal junction *(Figure 4)*. After tumescent anesthesia, the vein is ablated in a retrograde fashion. The postablation procedures are similar for both techniques.
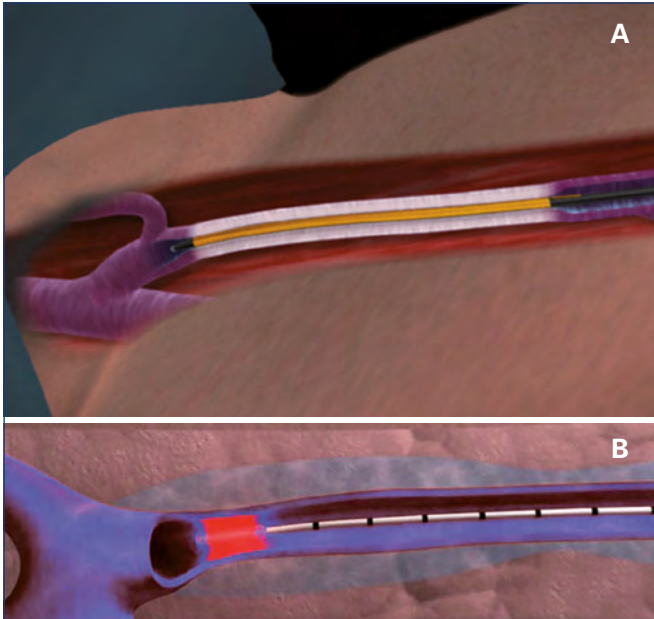
Figure 4. Positioning of the ClosureFAST catheter.

Panel A. The ClosureFAST catheter is positioned 2 cm below the saphenofemoral junction at the beginning of the procedure before generator activation. Panel B. The laser fiber catheter is positioned 2 cm below the saphenofemoral junction at the beginning of the procedure before activation. The veins are colored blue.

Endovenous steam ablation

In 2006, Milleret et al introduced steam as a cheaper alternative to laser and radiofrequency ablation.[15] The principle consists of injecting pulses of water vapor at 120°C in the vein to be ablated, with each pulse delivering 60 J of energy into the lumen. Steam is injected under pressure, whereby the first pulse dislodges the blood and the subsequent ones heat the vein wall. A 5F gauge stainless steel catheter is used because it is flexible enough to navigate through the tortuosity without using a guide wire. Two lateral holes close to the tip eject the steam, avoiding the risk of heating deep veins when heating the junctions.

A comparative animal study by Thomis et al compared steam with either ClosureFAST radiofrequency or a 1470 nm TULIP fiber laser. The three methods generated comparable results regarding scores for low perivenous tissue destruction and high vein wall destruction.[13]

In a pilot study by van den Bos et al, 11 out of the 19 veins treated were completely obliterated at 6 months, with a partial reopening in the other veins. However, the energy delivered was too low, 1 pulse/cm instead of the 2 to 4 pulses/cm that is advised by the manufacturers of the technique.[14] In a series of 75 patients, the complications included a thrombus protrusion in the femoral vein, an ecchymosis at the entry site in 1 patient, and moderate pain lasting 8 days in 6 patients. Subsequently, a randomized controlled trial was designed and it is still ongoing.[15]

Endovenous microwave ablation

After ligation of the saphenofemoral junction, the microwave treating wire is inserted into the great saphenous vein until the medial aspect of the ankle and is guided by the illuminated tip of the wire. The treating wire is withdrawn from distal to proximal at 2 to 4 mm/s, delivering 80 J/cm of energy. In 16.4% of patients, the treating wire could not be passed to the ankle; therefore, it was inserted in the great saphenous vein at a puncture in the ankle and the vein ablation was conducted from groin to ankle. In the same session, all superficial varicose veins and perforators are ablated using short-wire power (10 to 15 W) under ultrasound guidance.[16]

Complications of endovenous thermal ablation

In a review analyzing randomized controlled trials conducted on open surgery (radiofrequency ablation 317 patients), endovenous laser ablation (1057 patients), and open surgery (975 patients), the short-term complications included venous thromboembolism, wound infection, and paresthesia.[17] There was a significantly higher rate of wound infection for open surgery (2.3%; 95% CI, 1.3%-3.1%) vs endovenous laser ablation (0.5%; 95% CI, 0.3%-1.3%; P=0.006), but not between open surgery and radiofrequency ablation (1.5%; 95% CI, 0.4%-3.0%; P=0.094). The paresthesia rate was significantly lower with endovenous laser ablation (3.8%; 95% CI, 2.4%-4.5%) compared with radiofrequency ablation (5.2%; 95% CI, 3.1%-7.9%; P<0.001) and open surgery (7.4%; 95% CI, 5.3%-8.3%; P<0.001). The rate of thrombophlebitis was significantly lower for open surgery (3.0%; 95% CI, 2.9%-4.0%) compared with both radiofrequency ablation (5.5%; 95% CI, 3.0%-7.8%; P=0.003) and endovenous laser ablation (5.6%; 95% CI, 4.2%-7.0%; P=0.003). Thermal skin burns occurred with equal frequency between radiofrequency ablation and endovenous laser ablation.[17]

A review of radiofrequency ablation complications has been reported and this method has been compared with those of other operative procedures. Early complications include pain, phlebitis (7% to 9.6%), arteriovenous fistula (0.15%), endovenous heat-induced thrombosis (EHIT), deep vein thrombosis (<0.01%), lidocaine toxicity, wound problems (6% to 8%), and skin burns (0.5%). Late complications are mostly transient and may include skin pigmentation (6% to 19%) and nerve damage (4% to 20%).[18,19] Complications from endovenous laser ablation have also been compiled and include phlebitis (1.87%), skin burns (0.46%),

nerve injury (3.08%), arteriovenous fistula (0.15%), endovenous heat-induced thrombosis, and deep venous thrombosis (0.27%).[19]

Only one multicenter trial has reported the outcomes of endovenous steam ablation (n=117). Postprocedural pain was lower in endovenous steam ablation compared with endovenous laser ablation. Other outcomes included thrombophlebitis (9.2%), nerve injury (0.9%), and hyperpigmentation (4.6%), but no deep vein thrombosis or skin burns were identified.[20] Complications after endovenous microwave ablation have been reported in a single-center study, where endovenous microwave ablation was responsible for skin burns related to ablation of subcutaneous tributaries (10.2%).[16]

Chemical ablation

Sclerotherapy

Sclerotherapy refers to the introduction of a foreign substance into the lumen of a venous vessel to damage the venous wall and occlude the vessel. Liquid sclerotherapy has been used primarily for obliteration of spider veins. However, interest in using sclerotherapy for telangiectasia and varicose veins significantly increased in 1995 when Cabrera et al reported that foam, prepared by mixing gas with the detergent polidocanol, was effective for obstruction of larger veins.[21] The use of ultrasound-guided foam sclerotherapy has rapidly spread for the treatment of primary and recurrent varicose veins, including the great saphenous vein, small saphenous vein, saphenous tributaries, and perforating veins.

Sclerosing agents

The mechanism of action for sclerosing agents includes destruction of venous endothelial cells, exposure of subendothelial collagen fibers, and ultimately, the formation of a fibrotic obstruction. Delivery of the solution as a foam prolongs the contact time and amplifies the effect of the chemical substance. For producing endothelial injury, sclerosing solutions can be classified into three categories: detergent, osmotic, or chemical irritant.

In Europe, approved agents for sclerotherapy include sodium tetradecyl sulfate, polidocanol, morrhuate sodium, hypertonic saline, and glycerin.

- Sodium tetradecyl sulfate is a detergent that destroys the endothelium by denaturation of the cell surface proteins. The solution is safe and painless when injected. When the solution is injected at higher concentrations, extravasation may result in tissue necrosis. Hyperpigmentation, matting, and allergic reactions have been described, but rarely occurred. Generating foam with a sodium tetradecyl sulfate agent is easy.
- Polidocanol is another detergent that is safe and painless when injected and has a low risk of tissue necrosis when used at low concentrations. It may cause hyperpigmentation, but has a very low rate of allergic or anaphylactic reactions. There is a consensus that polidocanol has fewer overall complications compared with sodium tetradecyl sulfate.

- Sodium morrhuate is a detergent that is used less frequently due to a relatively higher incidence of skin necrosis observed with extravasation and a higher risk of anaphylactic reactions within a few minutes after injection.
- Glycerin is a chemical irritant that destroys the cell surface proteins by affecting chemical bonds. Chromated glycerin is frequently used as a solution of glycerin, sterile water, and benzyl alcohol. Chromated glycerin is safe and rarely leads to tissue necrosis, hyperpigmentation, or allergies, but frequently there is local pain at the injection site. This treatment is particularly suitable for treating small veins or telangiectasia.
- Hypertonic saline, an osmotic agent, is a weak sclerosing agent that causes dehydration of endothelial cells through osmosis, which leads to endothelial cell death. Burning pain is frequent during injection. Extravasation may cause skin ulcers and tissue necrosis.

Liquid sclerotherapy

Liquid sclerotherapy is currently used for treating reticular veins and telangiectasia.

Foam sclerotherapy

Due to the enhanced sclerosing properties of foam, ultrasound-guided foam sclerotherapy has been shown to be more effective than liquid sclerotherapy, Tessari et al used a three-way stopcock connected to two syringes to produce foam and they developed the most popular technique used today.[22] Other techniques for producing foam involve a two-way female-to-female connector.

Experts recommend a ratio of 1 part sodium tetradecyl sulfate or polidocanol to 4 or 5 parts air. Mixing the drug with air using the two syringes and pushing the mixture from one syringe into the other 20 times results in an approximate bubble size of <100 μm. Coleridge Smith advises puncturing the veins in supine patients and then elevating the limb 30 degrees to inject the foam.[23] Ultrasonography is used to monitor the movement of foam in the veins. The saphenous vein is injected first, followed by varicose and perforating veins, if indicated. A maximum of 10 mL of foam is injected during one session. The procedure is completed by placing a short-stretch bandage or a 30 to 40 mm Hg graduated compression stocking on the limb. Most experts recommend 1 to 2 weeks of compression.

Severe complications of ultrasound-guided foam sclerotherapy comprise anaphylaxis (extremely rare), large tissue necrosis (extremely rare), stroke and transient ischemic attack (extremely rare), distal deep venous thrombosis (very rare), pulmonary embolism (extremely rare), and motor nerve injury (extremely rare). Benign complications are visual disturbances (uncommon), headaches and migraines (uncommon), sensory nerve injury (rare), chest tightness (very rare), dry cough (very rare), superficial thrombophlebitis (unclear), skin reaction (very rare), matting (common), residual pigmentation (common), minimal skin necrosis (very rare), and embolia cutis medicamentosa (very rare).

The complications are listed in the European guidelines for sclerotherapy in chronic venous disorders, along with recommendations to avoid and manage these complications. Ultrasound-guided foam sclerotherapy of the saphenous vein is the least invasive of the endovenous ablation techniques. In 2008, the European

Consensus Meeting on Foam Sclerotherapy reported that foam was an effective, safe, and minimally invasive endovenous treatment for varicose veins with a low rate of complications.[24] The most complete book on sclerotherapy was written by a team of editors in 2007.[25]

Cyanoacrylate glue ablation

A new nonablative procedure that intravenously delivers a cyanoacrylate adhesive mixture has been developed to improve some of the limitations of radiofrequency ablation, endovenous laser ablation, and sclerotherapy ablation. Upon intravascular injection, the cyanoacrylate adhesive rapidly solidifies via a polymerization reaction and results in an inflammatory reaction in the vein wall.

The disposable Sapheon Closure System includes 4 mL of Sapheon Cyanoacrylate Adhesive (SCA) and a Sapheon delivery system *(Figure 5 and 6)*. The Sapheon delivery system consists of a 7F-introducer sheath/dilator, a 5F-delivery catheter, a 3 mL syringe, and a dispenser gun. The hydrophobic 5F-delivery catheter has a novel configuration with air-filled microchannels to enhance sonographic visibility. The dispenser gun will deliver 0.08 to 0.16 mL of SCA with each trigger pull. Access to the great saphenous vein is achieved by applying the Seldinger technique, which uses a standard micropuncture kit under ultrasound localization. The Sapheon introducer sheath and dilator is advanced to the saphenofemoral junction over a 0.035 J guide wire.[26] The cyanoacrylate adhesive is extracted from its glass vial and loaded into a syringe, which is then attached to the 5F delivery catheter. The combined syringe and catheter are connected to a dispenser gun. The catheter is then primed by advancing the glue with the dispenser gun to within 3 cm of the catheter tip. To prevent thrombus extension through the saphenofemoral junction, the hydrophobic delivery catheter is placed approximately 5 cm below the saphenofemoral junction. The saphenofemoral junction is manually compressed with the ultrasound transducer and the proprietary adhesive is delivered using the Sapheon delivery system using two injections at 1 cm intervals. Compression of the saphenofemoral junction and the delivery site is maintained for 3 minutes. The adhesive is delivered at 3 cm intervals through the remainder of the target vein using 30 seconds of compression for each subsequent delivery of adhesive *(Figure 6)*. The last injection site is 2 to 4 cm from the entrance site to prevent the glue from migrating outside the vein. After venous closure is confirmed by ultrasound imaging, the catheter is removed and compression is applied to the catheter entry site until hemostasis is achieved. A single adhesive bandage is applied; neither compression stockings nor compression bandages are used. This protocol has been described in details in two articles.[26,27] Postoperative complications were minimal.

Figure 5. Sapheon kit that includes the Sapheon delivery system and the Sapheon cyanoacrylate adhesive flask.
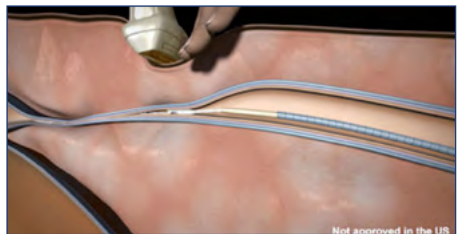

Figure 6. Compression of the treated vein using an ultrasound transducer above the catheter and injected glue.

Almeida et al reported a series of 38 patients treated for great saphenous vein incompetence. Postoperative side effects included a thread-like thrombus or glue extension across the saphenofemoral junction (21.1%), which resolved at 3 months, transient thrombophlebitis (16%), and hyperpigmentation (2.4%).[28] In another series including 43 great saphenous veins and 22 small saphenous veins, thrombophlebitis of the great saphenous vein occurred 4 times.[26] The primary potential advantage with this new technique is that it does not require tumescent anesthesia and patients do not need postoperative compression stockings.

Mechanochemical ablation

Recently, a new hybrid mechanochemical device (ClariVein) has been developed. Mechanochemical endovenous ablation (MOCA) achieves venous occlusion by utilizing a wire within the lumen of the vein that rotates at 3500 rpm, which abrades the intima and causes venospasms, thereby increasing the efficacy of the sclerosant *(Figures 7 and 8)*. A liquid sclerosant (sodium tetradecyl sulfate or polidocanol) is concomitantly infused through an opening close to the distal end of the catheter near the rotating wire. These two modalities—mechanical and chemical—achieve venous occlusion results equal to endothermal methods. The system includes an infusion catheter, motor drive, stopcock, and syringe. The dispersion wire extends through the catheter lumen and it is connected to an interface cartridge unit for connection to the 9V DC battery of the motorized handle unit on the proximal end, which controls wire rotation. The handle unit also provides a grip and syringe holder to facilitate physician-controlled infusion. The wire and the catheter sheath are inserted percutaneously into the vein under site anesthesia while the patient is in a reversed Trendelenburg position. The catheter sheath is retracted to expose the wire tip, which is positioned 2 cm from the saphenofemoral junction. The patient is then rotated into a flat position for the remainder of the procedure. The catheter motor is turned on and the catheter is pulled down the vein at a rate of approximately 1 to 2 mm/second, while the wire rotates and the sclerosing agent is infused. After removal of the catheter, occlusion of the great saphenous vein and patency of the common femoral vein is checked by duplex ultrasound.
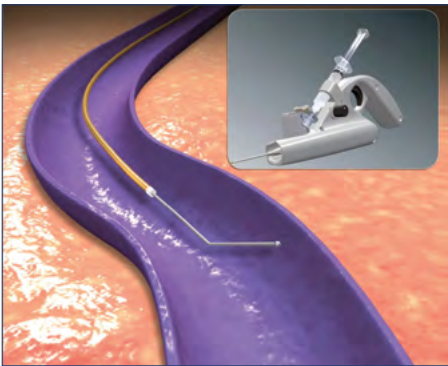

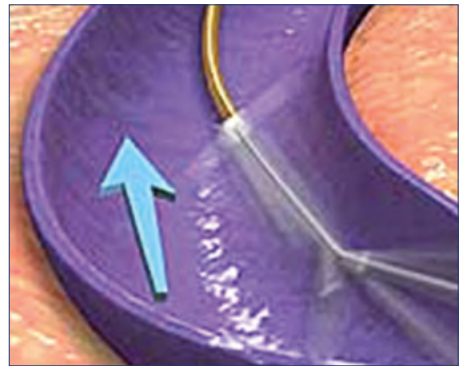*Figure 7. The vein lumen was catheterized using the ClariVein rotating wire.*

Figure 8. The ClariVein rotating wire abrades the vein wall, while the sclerosing agent is infused through the catheter opening.

The advantages of this hybrid system are claimed to be standard percutaneous access, endovenous treatment, local anesthesia only (without the need for tumescent anesthesia), and short procedure time. Since the system does not use thermal energy, the potential for nerve damage is minimized. Compression is applied for 2 weeks without restricting the patient's activity.[29]

In a small series of 25 patients presenting with great saphenous vein incompetence, minor postoperative complications were identified, including localized ecchymosis at the puncture site in 9 patients and transient thrombophlebitis of distal tributaries in 4 patients.[30] In a series of 50 patients presenting with small saphenous vein incompetence, minor postoperative complications were identified, including localized ecchymosis induration around the puncture site (12%) and transient thrombophlebitis of the treated vein (14%).[31]

Pelvic and ovarian vein embolization

When varicose veins are fed by incompetent pelvic and ovarian veins through the pelvic floor, which may or may not be related to left renal or iliac vein compression, embolization of the refluxive veins by coils and sclerosing agents is a minimally invasive method. Nevertheless, when reflux is related to iliac vein compression iliac stenting, another noninvasive technique, is the first-line treatment.[32,33]

Outcomes after operative treatment

Randomized controlled trials (RCTs) are very good tools for comparing the results of the various operative treatments for varicose veins.[34,35] Yet, before drawing definitive conclusions on any of these procedures, an accurate publication analysis is mandatory as RCTs often contain hard-to-identify bias. For example, the short-term results of a procedure greatly depend on the type of anesthesia performed during varicose vein ablation (local tumescent anesthesia or general anesthesia).[36] In the absence of RCTs for evaluating a procedure, case series are considered even though they provide a weaker level of evidence. Well-designed meta-analyses can provide valuable information for clinicians. By combining RCTs, meta-analyses increase the sample size, and thus, the power to study the results of a given procedure. Study outcomes are usually divided into the following 3 categories: (i) postoperative outcomes (<1 month); (ii) short- to mid-term outcomes (1 month to 3 years); and (iii) long-term outcomes (>3 years for RCTs and >5 years for case series. Nevertheless, this review's outcome analysis has been divided into two parts: (i) postoperative and mid-term outcomes and (ii) long-term outcomes.

Postoperative and mid-term outcomes

Open surgery

Classic open surgery has been compared with conservative treatment both in C_2 and C_5-C_6 patients *(Tables I.1 and I.2).*[37-48] In addition, classic open surgery has been compared with open surgery variants

	Operative procedures	Reference(s)
1	Classic open surgery vs Conservative treatment	Michaels et al,[37] 2006 Michaels et al,[38] 2006 Ratcliffe et al,[39] 2006 Sell et al,[40] 2014
2	Classic open surgery ± SEPS or laser ablation + compression therapy vs Isolated compression therapy in C_5-C_6 or C_6 patients	Barwell et al,[41] 2004 Guest et al,[42] 2003 Gohel et al,[43] 2005 van Gent et al,[44] 2006 Gohel et al,[45] 2007 Zamboni et al,[46] 2003 Zamboni et al,[47] 2004 Viarengo et al,[48] 2007
3	Classic open surgery vs Cryostripping	Menyhei et al,[49] 2008 Klem et al,[50] 2009
4	Classic open surgery with various types of tributary phlebectomy	Aremu et al,[51] 2004 Scavée et al,[52] 2003 Ray-Chaudury et al,[53] 2003 Chetter et al,[54] 2006 Krasznai et al,[55] 2015
5	Classic open surgery: partial vs complete stripping	Holme et al,[56] 1990
	Classic open surgery: HL comparing two skin closure techniques	Corder et al,[57] 1991
	Classic open surgery: HL + tributary phlebectomy vs Isolated HL	Dwerryhouse et al,[58] 1999
	Classic open surgery with and without a tourniquet	Sykes et al,[59] 2000
	Classic open surgery with SFJ flush ligation + tributary phlebectomy vs SFJ distal ligation + tributary phlebectomy	Belcaro et al,[60] 2002
	Classic open surgery with saphenous stripping (Babcock) vs Pin stripping (Oesch)	Buttler et al,[61] 2002

Table I. (page 17 to page 23)

	Operative procedures	Reference(s)
5	Classic open surgery under general + local anesthesia: lidocaine + adrenaline vs Saline solution	Nisar et al,[62] 2006
	Classic open surgery with saphenous stripping (Babcock) vs Invaginated stripping	Scheltinga et al,[63] 2007
	Classic open surgery with HL + stripping + tributary phlebectomy vs Idem + SEPS	Kianifard et al,[64] 2007
	Redo open surgery with SFJ ligation vs Redo SFJ ligation + polytetrafluoroethylene patch insertion in recurrent great saphenous varicose veins	Winterborn et al,[65] 2007
	Chemical ablation (UGFS) + HL vs HL + stripping	Abela et al,[66] 2008
	Flush SFJ ligation vs Standard transfixion SFJ ligation	Winterborn et al,[67] 2008
	HL + stripping + tributary phlebectomy + antibiotic prophylaxis vs Idem <u>without</u> antibiotic prophylaxis	Mekako et al,[68] 2010
	Classic open surgery with HL of the SFJ vs Idem <u>without</u> high SFJ ligation	Casoni et al,[4] 2013
	HL vs HL + fascia cribriformis suture vs HL with inverting suture of the stump	Haas et al,[69] 2005
6	Classic open surgery vs CHIVA	Carandina et al,[70] 2008 Parés et al,[71] 2010

	Operative procedures	Reference(s)
7	Classic open surgery vs RFA	Hinchliffe et al,[72] 2006 Kianifard et al,[73] 2006 Lurie et al,[74] 2003 Lurie et al,[75] 2005 Rautio et al,[76] 2002 Perälä et al,[77] 2005 Stötter et al,[78] 2006 Subromania et al,[79] 2010 Elkaffas et al,[80] 2011
8	Classic open surgery vs EVLA	de Medeiros et al,[81] 2005 Vuylstecke et al,[82] 2006 Lin et al,[83] 2007 Rasmussen et al,[84] 2007 Darwood et al,[85] 2008 Kalteis et al,[86] 2008 Theivacumar et al,[87] 2009 Christenson et al,[88] 2010 Pronk et al,[89] 2010 Rasmussen et al,[90] 2010 Carradice et al,[91] 2011 Carradice et al,[92] 2011 Rass et al,[93] 2012 Rasmussen et al,[94] 2013 Flessenkämpfer et al,[95] 2013 Samuel et al,[96] 2013 Roopram et al,[97] 2013
9	Classic open surgery vs Endovenous thermal ablation (EVLA, RFA)	Dzieciuchowicz et al,[98] 2014

	Operative procedures	Reference(s)
10	Liquid chemical ablation vs Classic open surgery	Einarsson et al,[99] 1993
	Liquid chemical ablation + HL vs Classic open surgery	Rutgers et al,[100] 1994
	Liquid chemical ablation vs Classic open surgery + liquid chemical ablation vs Classic open surgery	Belcaro et al,[101] 2000
	Liquid and foam chemical ablation vs Various open surgery procedures	Belcaro et al,[102] 2003
	Phlebectomy vs Liquid chemical ablation	de Roos et al,[103] 2003
	Chemical ablation + HL vs Classic open surgery (HL + stripping)	Abela et al,[66] 2008 Bountouroglou et al,[104] 2006 Liu et al,[105] 2011 Kalodiki et al,[106] 2012
	Chemical ablation (UGFS) vs Classic open surgery (HL + stripping)	Figueiredo et al,[107] 2009 Shadid et al,[108] 2012
	Chemical ablation (liquid or foam) vs HL or HL + stripping or phlebectomy	Wright et al,[109] 2006
11	Classic open surgery vs EVLA vs UGFS	Biemans et al,[110] 2013 Brittenden et al,[111] 2014 Tassie et al,[112] 2014

	Operative procedures	Reference(s)
12	Classic open surgery vs EVLA vs UGFS vs RFA	Rasmussen et al,[113] 2011 Rasmussen et al,[114] 2013
13	Classic open surgery vs Endovenous steam ablation	WoźniakW et al,[115] 2015
14	HL + stripping + tributary phlebectomy+ perforators ligation vs HL + EMA of the GSV + EMA tributary phlebectomy + EMA perforators ablation	Yang et al,[16] 2013
15	Classic open surgery (HL + stripping) vs HL + tributary phlebectomy ± perforator ligation	Campanello et al,[116] 1996 Hammarsten et al,[117] 1990 Hammarsten et al,[118] 1993 Winterborn et al,[119] 2004
16	RFA vs EVLA	Almeida et al,[120] 2009 Shepherd et al,[121] 2010 Gale et al,[122] 2010 Goode et al,[123] 2010 Nordon et al,[124] 2011
17	RFA vs Invagination stripping vs Cryostripping	Stötter et al,[78] 2006
18	RFA completed with deleted or synchronized ambulatory incompetent tributary avulsion	Lane et al,[125] 2015
19	EVLA vs Endovenous steam ablation	van der Bos et al,[20] 2014

	Operative procedures	Reference(s)
20	EVLA with different wavelengths	Kabnick et al,[133] 2006
	HL + EVLA vs EVLA <u>without</u> HL	Disselhoff et al,[134] 2008 Disselhoff et al,[135] 2011
	EVLA of above-knee GSV vs Above- and below-knee GSV ablation	Theivacumar et al,[136] 2008
	EVLA with and without nitroglycerin ointment	Hogue et al,[137] 2008
	EVLA using 980 nm bare-tip fiber vs EVLA using 1470 nm radial fiber	Doganci et al,[125] 2010
	EVLA using 1470 nm radial fiber comparing warm and cold tumescence anesthesia	Pannier et al,[138] 2010 Dumantepe et al,[139] 2015
	EVLA using 980 nm vs 1500 nm diode	Vuylsteke et al,[140] 2011
	EVLA using a bare fiber vs EVLA using a tulip fiber	Vuylsteke et al,[12] 2012
	EVLA with 2- vs 7-day postoperative compression therapy	Bakker et al,[141] 2013
	EVLA using 12 W laser power with intermittent withdrawal vs 14 W laser power with continuous withdrawal	Samuel et al,[142] 2013
21	Sclerotherapy using polidocanol vs Saline solution	Kahle et al,[143] 2004

	Operative procedures	Reference(s)
22	Liquid sclerotherapy vs Foam sclerotherapy	Hamel-Desnos et al,[144] 2003 Yamaki et al,[145] 2004 Alòs et al,[146] 2006 Ouvry et al,[147] 2008 Rabe et al,[148] 2008
23	Sclerosing agent at various doses and concentrations	Hamel-Desnos et al,[149] 2005 Ceulen et al,[150] 2007 Hamel-Desnos et al,[151] 2007 Blaise et al,[152] 2010
24	Different compression therapy regimens after foam sclerotherapy	O'Hare et al,[153] 2010
	Foam sclerotherapy with and without compression therapy	Hamel-Desnos et al,[154] 2010
	In vivo biological effects of foam sclerotherapy	Hamel-Desnos et al,[155] 2011
25	EVLA + phlebectomy vs UGFS	Lattimer et al,[130] 2012 Lattimer et al,[131] 2012 Lattimer et al,[132] 2013
26	Visual foam sclerotherapy alone vs Visual + UGFS	Yamaki et al,[156] 2012
27	Foam sclerotherapy in thrombophilic patients in combination with thromboprophylaxis: low-molecular-weight heparin vs warfarin	Hamel-Desnos et al,[157] 2009
28	Ulcer healing and ulcer recurrence according to the presence or absence of incompetent perforators after SEPS	van Gent et al,[182] 2015
29	EVLA vs Cryostripping	Disselhoff et al,[127] 2008 Disselhoff et al,[128] 2008 Disselhoff et al,[129] 2009

Table I. Randomized controlled trials, case series, and meta-analyses comparing operative procedures for the treatment of primary superficial vein incompetence.

For more information on the trials, please go to www.phlebolymphology.org.

Abbreviations: CHIVA, Cure Hémodynamique de l'Insuffisance Veineuse en Ambulatoire (Conservative ambulatory HemodynamIc management of VAricose veins); EMA, endovenous microwave ablation; EVLA, endovenous laser ablation; GSV, great saphenous vein; HL, high ligation; RFA, radiofrequency ablation; SEPS, subfascial endoscopic perforator surgery; SFJ, saphenofemoral junction; UGFS, ultrasound-guided foam sclerotherapy.

(Tables I.3 and I.4), such as cryostripping[49,50] and tributary-powered phlebectomy[51-55]—techniques that are only rarely used in current practice. Some RCTs (Table I.5)[4,56-69] provide interesting information on how cryostripping influences nerve damage,[56,59,63] the short- and long-term outcomes according to the procedure used,[58,64,67] the results following saphenofemoral junction ablation and ligation[4,60,69] or associated perforator ablation.[61] The RCTs comparing classic open surgery with other ablative procedures are more interesting and are shown in Table I.6 to I.15.[70-120] Additionally, the CHIVA method is performed under local anesthesia when other open surgery techniques need spinal or general anesthesia, and as a result, CHIVA shortens the length of the hospital stay (Table I.6).[70-71]

All RCTs that compared the short-term results of classic open surgery with radiofrequency ablation, endovenous laser ablation, endovenous steam ablation,[115] endovenous microwave ablation, ultrasound-guided foam sclerotherapy (UGFS), and high ligation with tributary phlebectomy concluded that both endovenous procedures and high ligation with tributary phlebectomy are less painful than classic open surgery and these procedures shorten the time required before returning to normal activity. Sensory impairment and ecchymosis are less severe with endovenous microwave ablation than open surgery, even though endovenous microwave ablation causes skin burns, 10% of which are related to slow probe withdrawal or using energy that is too high (Table I.14).[16] However, when modern open surgery is performed under local anesthesia (unfortunately by very few teams), it is as effective postoperatively as any endovenous procedure.

Endovenous procedures

Endovenous procedures have been widely studied and compared with open surgery and other endovenous procedures.

Thermal ablation

Radiofrequency ablation. Radiofrequency ablation has been compared with open surgery, cryostripping, invagination stripping, endovenous laser ablation, and ultrasound-guided foam sclerotherapy (Table I.7, I.12, I.16, and I.17).[72-80,113,114,120-124] Studies of endovenous laser ablation using bare fibers vs radiofrequency ablation favored the latter since it is less painful and results in less ecchymosis. However, it is now acknowledged that radial fibers, which are currently used, provide better postoperative results than bare fibers.[126] No differences in efficacy and undesirable effects were observed between radiofrequency ablation and ultrasound-guided foam sclerotherapy in a 4-arm study.[113,114] At a 1-year follow-up, redo operations were less frequent after radiofrequency ablation compared with deleted or synchronized ambulatory incompetent tributary avulsion (Table I.18).[126]

Endovenous laser ablation. Treating varicose veins with endovenous laser ablation is a safe procedure in patients with active ulcers. Ulcers healed faster after endovenous laser ablation than in patients undergoing compression therapy alone and no ulcer recurrence occurred during a 1-year period posttreament.[48] Endovenous laser ablation has been compared with open surgery, cryostripping, invagination stripping, endovenous laser ablation, and ultrasound-guided foam sclerotherapy (Table I.8, I.11, I.12, I.16,

*and I.*19).[20,81-97,110-114,120-124] endovenous laser ablation and cryostripping *Table I.29*[127-129] were similarly effective in patients with varicose veins,[127,128] and endovenous laser ablation had a similar, but slightly higher, cost.[129]

When comparing ultrasound-guided foam sclerotherapy and endovenous laser ablation *(Table I.11 and I.25),*[110-112,130-132] no differences at 3 months[130,131] were observed for clinical results or vein obliteration, but ultrasound-guided foam sclerotherapy outperformed endovenous laser ablation in cost, treatment duration, postoperative pain reduction, and recovery. At 15 months,[132] there were no differences in clinical results, but vein occlusion was higher with endovenous laser ablation. At a 1-year follow-up, Biemans et al found no difference between the endovenous laser ablation and ultrasound-guided foam sclerotherapy in complications and clinical results, but ultrasound-guided foam sclerotherapy resulted in lower occlusion rates.[110] Brittenden et al showed similar clinical efficacy between ultrasound-guided foam sclerotherapy and endovenous laser ablation, but endovenous laser ablation had fewer complications and ultrasound-guided foam sclerotherapy had lower ablation rates at both 6 weeks and 6 months posttreatment.[111] Tassie et al showed that endovenous laser ablation has the highest probability of being cost-effective compared with classic open surgery and ultrasound-guided foam sclerotherapy.[112]

The 1-year treatment success of high-dose endovenous laser ablation was not inferior to that of endovenous steam ablation. Several secondary outcomes (eg, painful legs, patients' satisfaction, duration of analgesia, and limitations in daily life) were in favor of endovenous steam ablation (*P*<0.001).[20]

Data from ten RCTs on endovenous laser ablation variants *(Table I.20)*[12,125,133-142] show that: (i) below-knee endovenous laser ablation was not associated with saphenous nerve injury[136]; (ii) lower postoperative pain and better Venous Clinical Severity Scores (VCSS) were obtained with radial fibers compared with bare fibers[125] or tulip fibers[12]; (iii) cold tumescent anesthesia had fewer side effects and a reduction in analgesic intake than warm tumescent anesthesia[138,139]; and (iv) symptom intensity was lower and quality of life better when compression was applied for 2 to 7 days posttreatment.[141]

Chemical ablation

Sclerotherapy. Postoperative, short-term, and mid-term results are difficult to compare because many different protocols and outcome criteria were used *(Tables I.10 to I.12).*[99-114] RCTs on variants of sclerotherapy provide some data on postoperative course and short- or mid-term outcomes.[143-157] Foam sclerotherapy provides better results than liquid sclerotherapy *(Table I.22),*[143-148,156,157] and occlusion rates are similar when using either a 1% or 3% polidocanol foam solution *(Table I.23).*[149-152] The use of postoperative compression does not influence the percentage of patients with side effects after ultrasound-guided foam sclerotherapy *(Table I.25).*[153-155]

Glue. No RCTs evaluating glue vs other procedures have been conducted, but a case series has reported good results at a 2-year follow-up—occlusion rates were 92% and a significant improvement in VCSS was observed.[27]

Mechanochemical ablation

There are no RCTs for Clarivein®, but case series are available.[29-31] At a 6-month follow-up, the occlusion rate was 96% and the VCSS improved in a series of patient presenting with saphenous vein varices.[29] In the case series by Boersma et al on patients who underwent short saphenous vein ablation, the occlusion rate at 1 year was 94% and the VCSS improved.[31]

Long-term outcomes

Clinical parameters

PREVAIT

The term PREsence of Varices After operatIve Treatment (PREVAIT) was adopted in the VEIN-TERM transatlantic interdisciplinary consensus document.[158] PREVAIT is a frustrating problem for both the patients with varicose veins and the physicians who treat these varicose veins. Recurrent Varices After Surgery (REVAS) have been previously compared with classic open surgery.[159]

Severity scores

Three severity scores—VCSS, Venous Segmental Disease Score (VSDS), and Aberdeen Varicose Vein Questionnaire (AVVQ)—are used in the literature to assess treatment success rates. VCSS is a very good tool for evaluating the treatment of complicated varices, but it is less informative for uncomplicated C_2 patients.[160,161]

Generic and specific health-related quality of life questionnaires

Many health-related quality of life questionnaires have been used, including AVVQ, the Chronic Venous Insufficiency Quality of Life Questionnaire (CIVIQ), and the Specific Quality of Life and Outcome Response-Venous (SQOR-V), and the results have been compared with anatomic, hemodynamic, and clinical outcomes before and after operative treatment.[162] Patient-Reported Outcome Measures (PROMs) are new and very promising tools.[163]

Instrumental investigation measurements

These measurements rely on occlusion rates and hemodynamic function. It has been clearly identified that the correlation between clinical and investigational parameters is far from perfect.

Information provided by RCTs

Open surgery vs high ligation and tributary phlebectomy

These procedures were assessed in 3 RCTs with 4, 5, and 11 years of follow-up[58,65,116-119] and there were no differences in clinical outcomes. More redo surgery was performed in the group with high ligation and tributary phlebectomy, but preoperative and postoperative investigations were outdated in both groups.

Open surgery vs CHIVA

CHIVA was compared with classic open surgery in 2 RCTs with 5 and 10 years of follow-up *(Table I.6).*[70,71] Both RCTs favor CHIVA in terms of PREVAIT reduction, but bias was identified to weaken the authors' conclusions.

Open surgery vs radiofrequency ablation

Only one RCT comparing long-term outcomes (3-year) of open surgery with radiofrequency ablation is available and there was no difference in clinical results between the two groups,[77] but the Closure® catheter used was older and less efficient that the ClosureFAST® catheter.

Open surgery vs endovenous laser ablation

At a 5-year follow-up, a RCT comparing endovenous laser ablation with open surgery found no difference between the 2 groups in persistent reflux, PREVAIT, redo treatment, VCSS, and generic and specific health-related quality of life scores. In this trial, open surgery was minimally invasive and the endovenous laser ablation procedure used a bare fiber with a 980-nm diode laser and a stepwise laser withdrawal.[94]

Sclerotherapy vs various open surgery procedures

Belcaro et al reported two series with long-term follow-up data, but no conclusive results were obtained.[101,102] The RCT comparing ultrasound-guided foam sclerotherapy complemented by high ligation with open surgery at a 3- to 5-year follow-up was more informative,[103] showing that the treatment was equally effective in both groups, which was demonstrated by improvements in the VCSS, VSDS, and the generic health-related quality of life scores. At 5 years posttreatment, the AVVQ was significantly better in the open surgery group.[106]

Information provided by case series

Open surgery

The most documented outcomes are provided by classic open surgery, but most studies are retrospective. In a 34-year follow-up study, varicose veins were present in 77% of the lower limbs examined and most were symptomatic—58% were painful, 83% had a tired feeling, and 93% showed a reappearance of edema.[164] Two prospective studies concerning classic open surgery are available with a 5-year follow-up.[165,166] In both studies, patients were preoperatively investigated with duplex scanning and treated by high ligation, saphenous trunk stripping, and stab avulsion. In the Kostas et al series, 28 out of 100 patients had PREVAIT after 5 years, where the recurrent varices mainly resulted from neovascularization (8/28, 29%), new varicose veins as a consequence of disease progression (7/28, 25%), residual veins due to tactical errors (eg, failure to strip the great saphenous vein) (3/28, 11%), and complex patterns (10/28, 36%).[166]

In the van Rij series, 127 limbs (CEAP class C_2-C_6) were evaluated postoperatively by clinical examination, duplex scanning, and air plethysmography. At the clinical evaluation, recurrence of varicose veins was progressive from 3 months (13.7%) to 5 years (51.7%). In line with clinical changes, a progressive deterioration in venous function was measured by air plethysmography and reflux recurrence was assessed by duplex scanning.[165] These two studies showed that recurrence of varicose veins after surgery is common,

even in highly skilled centers. Even if the clinical condition of most affected limbs after surgery improved compared with before surgery, progression of the disease and neovascularization are responsible for more than half of the recurrences. Rigorous evaluation of patients and assiduous surgical techniques might reduce the recurrence resulting from technical and tactical failures.

Other procedures

A 5-year follow-up of a large series of patients treated with radiofrequency ablation using Closure plus catheter showed that vein occlusion and absence of reflux were present in 87.2% and 83.8% of patients, respectively. Symptoms, including pain, fatigue, and edema, significantly improved compared with preoperative status. The rate of PREVAIT progressed from 6 months (7.7%) to 5 years (27.4%).[167] Currently, no long-term results are available for Glue and Clarivein®.

Information provided by meta-analyses

Since 2009, six meta-analyses on operative treatment of primary varicose veins by open surgery, radiofrequency ablation, endovenous laser ablation, and ultrasound-guided foam sclerotherapy were identified—all produced similar conclusions.[168-173]

Final remarks concerning outcomes after operative treatment

The immediate postoperative course, including side effects, recovery time, and convalescence, is better in all other procedures compared with classic open surgery, but this point is questioned if modern and minimally aggressive open surgery is used. No differences in recurrence between classic open surgery compared with radiofrequency ablation and endovenous laser ablation are present at the mid- or long-term follow-up. PREVAIT is more frequent after ultrasound-guided foam sclerotherapy compared with other mentioned procedures, but PREVAIT can be easily and effectively treated with redo ultrasound-guided foam sclerotherapy.

Operative treatment indications

According to CEAP class and instrumental investigations

In patients with primary superficial reflux who are classified as C_2, indications for operative treatment rely on patient complaints, such as symptoms and cosmetics, and the extent and size of the varices. For patients in the C_3 to C_6 classes, operative treatment must be considered in all cases, except for the usual contraindications. However, in all clinical classes, nonvenous causes must be identified because venous symptoms are not pathognomonic and some signs, including edema and ulcers may be due to other etiologies. In the presence of axial deep primary reflux combined with primary varices, varicose veins must be treated first. However, we know that, in about 5% of patients, axial deep primary reflux is not corrected by varicose vein ablation[174] and its persistence is responsible for varices recurrence.[175,176]

When incompetent perforators are associated with primary varices, do they need to be treated in the same session? As no RCTs have compared the outcomes after varicose vein ablation with perforator

ablation + varices ablation, no evidence-based information is available. Nevertheless, we know that, in half of these patients, incompetent perforators are no longer identified after varices ablation.[177-179] To summarize, perforator ablation can be reserved for patients with persistent incompetent perforator vessels, abnormal hemodynamic parameters, or continued symptoms and/or signs (C_{4b}-C_6) after superficial ablative surgery.[179-181] Nevertheless, one RCT favors treating perforators in C_6 patients to prevent ulcer recurrence *(Table I.28)*.[182]

Operative treatment indication in PREVAIT patients

PREVAIT represents a particular situation in terms of indication. Managing patients with PREVAIT varies according to the clinical situation. Patients attending a routine follow-up, who are either asymptomatic or symptomatic, and possibly complaining of recurrences are managed differently than symptomatic patients who are complaining of cosmetic problems and presenting with complicated varices (C_3-C_6).[177] A consensus document agrees that ultrasound-guided foam sclerotherapy is the first-line treatment in almost all cases, except in patients presenting with varicose veins of the lower limbs that are fed by pelvic refluxive veins. The European guidelines for sclerotherapy assigned a Grade 1B to this procedure.[24] In the absence of RCTs, this recommendation is based on case series.[183,184]

Operative treatment choice

In practice, the choice of the procedure is frequently not made on evidence-based data, but on other factors, such as: (i) personal mastery of the different techniques—practitioners will favor the procedures they have mastered; (ii) coverage/reimbursement by the health services/health insurance, which varies from country to country; (iii) the patient's choice, which is influenced by possible postoperative problems, recovery time, time off work, the procedure that provides the easiest control of recurrences, and information from friends, literature, or the internet.

Guidelines

Recommendations from five guidelines are summarized in *Table II*. The guidelines of the Society for Vascular Surgery/American Venous Forum (SVS/AVF) were published in 2011.[185] Most recommendations remain valid, but are not fully applicable in Europe. The SVS/AVF guidelines were analyzed by a European team.[186] In 2013, the European Guide for Sclerotherapy was available, giving much information on sclerotherapy, including practical information.[24] In 2014, the European Venous Forum (EVF) and the International Union of Angiology (IUA) published a guidelines document on the management of chronic venous disorders.[187] The International guidelines on endovenous thermal ablation were published in 2015. This consensus document also provides many technical details.[188] The same year, the European Society for Vascular Surgery (ESVS) endorsed guidelines on the management of chronic venous disease.[189]

Most of these guidelines used the Guyatt grading scheme, which classifies recommendations as strong (grade 1) or weak (grade 2), according to the balance among benefits, risks, burdens, cost, and the degree of confidence in the estimates of benefits, risks, and burdens. It classifies quality of evidence as high (grade A), moderate (grade B), or low (grade C) according to factors, such as study design, consistency of the results, and directness of the evidence.[190] Only the ESVS guidelines used the European Society of Cardiology's

Operative procedure	SVS/AVF[186]	EVF/IUA[187]	
Classic open surgery	GSV 2B* SSV 1B*	2A*	
Modern surgery	NG	1B*	
CHIVA	2B*	NG	
ASVAL	2C*	NG	
EVLA or RFA	1B*	1A*	
Steam			
Clarivein®	NG	NG	
Glue	NG	NG	
UGFS	NG	1A*	
Thermal ablation vs UGFS (GSV)	1B*	NG	
Thermal ablation vs Surgery (GSV)	1B*	NG	
Surgery for PREVAIT	2C*	NG	
UGFS for PREVAIT	2C*	NG	
Endovenous thermal ablation for PREVAIT	2C*	NG	

Table II. Recommendations for operative procedures for the treatment of superficial refluxing veins from the recent guidelines.

*Guyatt's grading[190]

**Grading system of the European Society of Cardiology[191]

Abbreviations: ASVAL, Ablation Selective des Varices sous Anesthésie Locale (Ambulatory Selective Vein Ablation under Local anesthesia); AVF, American Venous Forum; CHIVA, Cure Hémodynamique de l'Insuffisance Veineuse en Ambulatoire (Conservative ambulatory HemodynamIc management of VAricose veins); EGS, European Guide for Sclerotherapy; EVLA, endovenous laser ablation; ESVS, European Society of Vascular Surgery; ETAV, Endovenous Thermal Ablation for Varicose Vein Disease; EVF, European Venous Forum; GSV, great saphenous vein; IUA, International Union of Angiology; IUP, International Union of Phlebology; NG, not graded; PREVAIT, PREsence of VArices after operatIve Treatment; SSV, small saphenous vein; SVS, Society of Vascular Surgery; UGFS, ultrasound-guided foam sclerotherapy.

ESVS[189]	ETAV/IUP[188]	EGS[24]
I B**		
NG	NG	NG
II b B**	NG	NG
II a B**	NG	NG
GSV IA** SSV IIaB**	1A*	NG
	1A*	
NG	NG	NG
NG	NG	NG
IIIA**	NG	1A-1C* according to vein diameter
IA**	NG	NG
IA**	NG	NG
NG	NG	NG
IIaB**	NG	NG
IIaB**	NG	NG

grading system. For each recommendation, the letter A, B, or C marks the level of current evidence. Weighing the level of evidence and expert opinion, every recommendation is subsequently marked as either class I, IIa, IIb, or III. The lower the class number, the more proven the efficacy and safety of a certain procedure.[191]

In 2013, the National Institute for Health and Care Excellence (NICE) published a document on varicose veins of the leg,[192] where the recommendations for people with confirmed varicose veins and truncal reflux were as follows:

- First, offer endothermal ablation (radiofrequency ablation for varicose veins [NICE interventional procedure guidance 8][193] and endovenous laser ablation for the long saphenous vein [NICE interventional procedure guidance 52][194]).
- If endothermal ablation is unsuitable, offer ultrasound-guided foam sclerotherapy (see NICE interventional procedure guidance 440[195]).
- If ultrasound-guided foam sclerotherapy is unsuitable, offer surgery.
- If incompetent varicose tributaries are to be treated, consider treating them at the same time.[192]

Conclusions

Currently, there are a number of surgical options for treating varicose veins, but there is no definitive system for identifying which people will benefit the most from interventional treatment and no established framework for the diagnosis and management of varicose veins. Conversely, perioperative investigations are well stated and described. In a review of the randomized controlled trials on the treatment of varicose veins, the authors concluded that there are many treatment options available for the ablation of varicose veins, not solely thermal ablation.[34,35] Operative treatment of primary varicose veins is currently performed using minimally invasive procedures, excluding spinal or general anesthesia. The problem is that the development of new procedures or devices is so rapid that when long-term outcomes are available, particularly for RCTs, the technique or material evaluated is frequently no longer used. Postoperative quality of life has improved, complications are far less frequent, and sick leave is shorter. The long-term frequency of PREVAIT is approximately the same for all techniques used, as long as the initial procedure has been correctly executed. To minimize the severity of PREVAIT, it is crucial to have regular patient follow-up and use ultrasound investigation to manage possible varices recurrence.

References

1. Franceschi C. *Theorie et Pratique de la Cure Conservatrice et Hémodynamique de l' Insuffisance Veineuse en Ambulatoire*. Precy-sous-Thil, France: Editions de l'Armancon; 1988.

2. Pittaluga P, Chastanet S, Rea B, Barbe R. Midterm results of the surgical treatment of varices by phlebectomy with conservation of a refluxing saphenous vein. *J Vasc Surg*. 2009;50:107-118.

3. Morrison C, Dalsing MC. Signs and symptoms of saphenous nerve injury after greater saphenous vein stripping: prevalence, severity, and relevance for modern practice. *J Vasc Surg*. 2003;38:886-890.

4. Casoni P, Lefebvre-Vilardebo M, Villa F, Corona P. Great saphenous vein surgery without high ligation of the saphenofemoral junction. *J Vasc Surg*. 2013;58:173-178.

5. Huang TW, Chen SL, Bai CH, Wu CH, Tam KW. The optimal duration of compression therapy following varicose vein surgery: a meta-analysis of randomized controlled trials. *Eur J Vasc Endovasc Surg*. 2013;45:397-402.

6. Mariani F, Marone EM, Gasbarro V, et al. Multicenter randomized trial comparing compression with elastic stocking versus bandage after surgery for varicose veins. *J Vasc Surg*. 2011;53:115-122.

7. Rudström H, Björck M, Bergqvist D. Iatrogenic vascular injuries in varicose vein surgery: a systematic review. *World J Surg*. 2007;31:228-233.

8. Sutton PA, El-Duhwaib Y, Dyer J, Guy AJ. The incidence of post operative venous thromboembolism in patients undergoing varicose vein surgery recorded in Hospital Episode Statistics. *Ann R Coll Surg Engl*. 2012;94:481-483.

9. van Rij AM, Chai J, Hill GB, Christie RA. Incidence of deep vein thrombosis after varicose vein surgery. *Br J Surg*. 2004;91:1582-1585.

10. Sam RC, Silverman SH, Bradbury AW. Nerve injuries and varicose vein surgery. *Eur J Vasc Endovasc Surg*. 2004;27:113-120.

11. Zamboni P, Franceschi C. *Principles of Venous Hemodynamics*. Hauppauge, NY: Nova Science Publishers; 2009.

12. Vuylsteke ME, Thomis S, Mahieu P, Mordon S, Fourneau I. Endovenous laser ablation of the great saphenous vein using a bare fibre versus a tulip fibre: a randomised clinical trial. *Eur J Vasc Endovasc Surg*. 2012;44:587-592.

13. Thomis S, Verbrugghe P, Milleret R, Verbeken E, Fourneau I, Herijgers P. Steam ablation versus radiofrequency and laser ablation: an in vivo histological comparative trial. *Eur J Vasc Endovasc Surg*. 2013;46:378-382.

14. van den Bos RR, Milleret R, Neumann M, Nijsten T. Proof-of-principle study of steam ablation as novel thermal therapy for saphenous varicose veins. *J Vasc Surg*. 2011;53:181-186.

15. Milleret R, Huot L, Nicolini P, et al. Great saphenous vein ablation with steam injection: results of a multicentre study. *Eur J Vasc Endovasc Surg*. 2013;45:391-396.

16. Yang L, Wang XP, Su WJ, Zhang Y, Wang Y. Randomized clinical trial of endovenous microwave ablation combined with high ligation versus conventional surgery for varicose veins. *Eur J Vasc Endovasc Surg*. 2013;46:473-479.

17. Dermody M, O'Donnel TF, Balk EM. Complications of endovenous ablation in randomized controlled trials. *J Vasc Surg Venous Lymphat Disord*. 2013;1:427-436.

18. Anwar MA, Lane TR, Davies AH, Franklin IJ. Complications of radiofrequency ablation of varicose veins. *Phlebology*. 2012;47(suppl 1):34-39.

19. Dexter D, Kabnick L, Berland T, et al. Complications of endovenous lasers. *Phlebology*. 2012;47(suppl 1):40-45.

20. van der Bos RR, Malskat WS, De Maeseneer MG, et al. Randomized clinical trial of endovenous laser ablation versus steam ablation (LAST trial) for great saphenous varicose veins. *Br J Surg*. 2014;101:1077-1083.

21. Cabrera J, Cabrera García-Olmedo JR. Nuevo método de esclerosis en las varices tronculares. *Patologia Vascular*. 1995;4:55-73.

22. Tessari L, Cavezzi A, Frullini A. Preliminary experience with a new sclerosing foam in the treatment of varicose veins. *Dermatol Surg*. 2001;27:58-60.

23. Coleridge Smith P. Chronic venous disease treated by ultrasound guided foam sclerotherapy. *Eur J Vasc Endovasc Surg*. 2006;32:577-583.

24. Rabe E, Breu FX, Cavezzi A, et al; Guideline Group. European guidelines for sclerotherapy in chronic venous disorders. *Phlebology*. 2014;29:338-354.

25. Goldman MP, Bergan JJ, Guex JJ. *Sclerotherapy: Treatment of Varicose and Telangiectatic Leg Veins*. 4th ed. Philadelphia, PA: Mosby Elsevier; 2007.

26. Lawson JK, Gauw S, van Vlijmen C, et al. Sapheon: the solution? *Phlebology*. 2013;28(suppl 1):2-9.

27. Almeida JI, Javier JJ, Mackay E, Bautista C, Proebstle TM. First human use of cyanoacrylate adhesive for treatment of saphenous vein incompetence. *J Vasc Surg Venous Lymphat Disord*. 2013;1:174-180.

28. Almeida JI, Javier JJ, Mackay EG, Bautista C, Cher DJ, Proebstle TM. Two-year follow-up of first human use of cyanoacrylate adhesive for treatment of saphenous vein incompetence. *Phlebology*. 2015;30:397-404.

29. Elias S, Lam YL, Wittens CH. Mechanochemical ablation: status and results. *Phlebology*. 2013;28(suppl 1):10-14.

30. van Eekeren RR, Boersma D, Elias S, et al. Endovenous mechanochemical ablation of great saphenous vein incompetence using the ClariVein® device: a safety study. *J Endovasc Ther*. 2011;18:328-334.

31. Boersma D, van Eekeren RR, Werson DA, van der Waal RI, Reijnen MM, de Vries JP. Mechanochemical endovenous ablation of small saphenous vein insufficiency using the ClariVein® device: one-year results of a prospective series. *Eur J Vasc Endovasc Surg*. 2013;45:299-303.

32. Bora A, Avcu S, Arslan H, Adali E, Bulut MD. The relation between pelvic varicose veins and lower extremity venous insufficiency in women with chronic pelvic pain. *JBR-BTR*. 2012;95:215-221.

33. Monedero JL, Ezpeleta SZ, Perrin M. Pelvic congestion syndrome can be treated operatively with good long-term results. *Phlebology*. 2012;27(suppl 1):65-73.

34. Eklöf B, Perrin M. Randomized controlled trials in the treatment of varicose veins. I. *Phlebolymphology*. 2011;18:196-208.

35. Perrin M, Eklöf B. Randomized controlled trials in the treatment of varicose veins. II. *Phlebolymphology*. 2012;19:92-99.

36. Thakur B, Shalhoub J, Hill AM, Gohel MS, Davies AH. Heterogeneity of reporting standards in randomised clinical trials of endovenous interventions for varicose veins. *Eur J Vasc Endovasc Surg*. 2010;40:528-533.

37. Michaels JA, Brazier JE, Campbell WB, MacIntyre JB, Palfreyman SJ, Ratcliffe J. Randomized clinical trial comparing surgery with conservative treatment for uncomplicated varicose veins. *Br J Surg*. 2006;93:175-181.

38. Michaels JA, Campbell WB, Brazier JE, et al. Randomised clinical trial, observational study and assessment of cost-effectiveness of the treatment of varicose veins (REACTIV trial). *Health Technol Assess*. 2006;10:1-196.

39. Ratcliffe J, Brazier JE, Campbell WB, Palfreyman SJ, MacIntyre JB, Michaels JA. Cost effectiveness analysis of surgery versus conservative treatment for uncomplicated varicose veins in a randomized control trial. *Br J Surg*. 2006;93:182-186.

40. Sell H, Vikatamaa P, Albäck A, et al. Compression therapy versus surgery in the treatment of patients with varicose veins: a RCT. *Eur J Vasc Endovasc Surg*. 2014;47:670-677.

41. Barwell JR, Davies CE, Deacon J, et al. Comparison of surgery and compression with compression alone in chronic venous ulceration (ESCHAR study): randomised controlled trial. *Lancet*. 2004;363:1854-1859.

42. Guest M, Smith JJ, Tripuraneni G, et al. Randomized clinical trial of varicose vein surgery with compression versus compression alone for the treatment of venous ulceration. *Phlebology*. 2003;18:130-136.

43. Gohel MS, Barwell JR, Earnshaw JJ, et al. Randomized clinical trial of compression plus surgery versus compression alone in chronic venous ulceration (ESCHAR study)—haemodynamic and anatomical changes. *Br J Surg*. 2005;92:291-297.

44. van Gent WB, Hop WC, van Praag MC, Mackaay AJ, de Boer EM, Wittens CH. Conservative versus surgical treatment of venous leg ulcers: a prospective, randomized, multicenter trial. *J Vasc Surg*. 2006;44:563-571.

45. Gohel MS, Barwell JR, Taylor M, et al. Long term results of compression therapy versus compression plus surgery in chronic venous ulceration (ESCHAR): a randomized controlled trial. *BMJ*. 2007;335:83.

46. Zamboni P, Cisno C, Marchetti F, et al. Minimally invasive surgical management of primary venous ulcers vs. compression treatment: a randomized clinical trial. *Eur J Vasc Endovasc Surg*. 2003;25:313-318.

47. Zamboni P, Cisno C, Marchetti F, et al. Haemodynamic CHIVA correction surgery versus compression for primary venous ulcers: first year results. *Phlebology*. 2004;19:28-34.

48. Viarengo LM, Potério-Filho J, Potério GM, Menezes FH, Meirelles GV. Endovenous laser treatment for varicose veins in patients with active ulcers: measurement of intravenous and perivenous temperatures during the procedure. *Dermatol Surg*. 2007;33:1234-1242.

49. Menyhei G, Gyevnàr Z, Aratá E, Kelemen O, Kollár L. Conventional stripping versus cryostripping: a prospective randomised trial to compare improvement in quality of life and complications. *Eur J Vasc Endovasc Surg*. 2008;35:218-223.

50. Klem TM, Schnater JM, Schütte PR, Hop W, van der Ham AC, Wittens CH. A randomized trial of cryostripping versus conventional stripping of the great saphenous vein. *J Vasc Surg*. 2009;49:403-409.

51. Aremu M, Mahendran B, Butcher W, et al. Prospective randomized controlled trial: conventional versus powered phlebectomy. *J Vasc Surg*. 2004;39:88-94.

52. Scavée V, Lesceu O, Theys S, Jamart J, Louagie Y, Schoevaerdts JC. Hook phlebectomy versus transilluminated powered phlebectomy for varicose veins surgery: early results. *Eur J Vasc Endovasc Surg*. 2003;25:473-475.

53. Ray-Chaudury S, Huq Z, Souter R, McWhinnie D. A randomized controlled trial comparing transilluminated powered phlebectomy with hook avulsions. An adjunct to day surgery? *J One-Day Surg*. 2003;13:24-27.

54. Chetter IC, Mylankal KJ, Hughes H, Fitridge R. Randomized clinical trial comparing multiple stab incision phlebectomy and transilluminated powered phlebectomy for varicose veins. *Br J Surg*. 2006;93:169-174.

55. Krasznai AG, Sigterman TA, Willems CE, et al. Prospective study of a single treatment strategy for local tumescent anesthesia in Muller phlebectomy. *Ann Vasc Surg*. 2015;29:586-593.

56. Holme JB, Skajaa K, Holme K. Incidence of lesions of the saphenous nerve after partial or complete stripping of the long saphenous vein. *Acta Chir Scand*. 1990;156:145-148.

57. Corder AP, Schache DJ, Farquharson SM, Tristram S. Wound infection following high saphenous ligation: a trial comparing two skin closure techniques: subcuticular polyglycolic acid and interrupted monofilament nylon mattress sutures. *J R Coll Surg Edinb*. 1991;36:100-102.

58. Dwerryhouse S, Davies B, Harradine K, Earnshaw JJ. Stripping the long saphenous vein reduces the rate of reoperation for recurrent varicose veins: five-year results of a randomized trial. *J Vasc Surg*. 1999;29:589-592.

59. Sykes TC, Brookes P, Hickey NC. A prospective randomised trial of tourniquet in varicose vein surgery. *Ann R Coll Surg Engl*. 2000;82:280-282.

60. Belcaro G, Nicolaides AN, Cesarone NM, et al. Flush ligation of the saphenofemoral junction vs simple distal ligation, 10 year, follow-up. The safe study. *Angéiologie*. 2002;54:19-23.

61. Butler CM, Scurr JH, Coleridge Smith PD. Prospective randomized trial comparing conventional (Babcock) stripping with inverting (Pin) stripping of the long saphenous vein. *Phebology*. 2002;17:59-63.

62. Nisar A, Shabbir J, Tubassam MA, et al. Local anaesthesic flush reduces postoperative pain and haematoma formation after great saphenous vein stripping—a randomised controlled trial. *Eur J Vasc Endovasc Surg*. 2006;31:325-331.

63. Scheltinga MR, Wijburg ER, Keulers BJ, de Kroon KE. Conventional versus invaginated stripping of the great saphenous vein: a randomized double-blind, controlled clinical trial. *World J Surg*. 2007;31:2236-2242.

64. Kianifard B, Holdstock J, Allen C, Smith C, Price B, Whiteley MS. Randomized clinical trial of the effect of adding subfascial endoscopic perforator surgery to standard great saphenous vein stripping. *Br J Surg*. 2007;94:1075-1080.

65. Winterborn RJ, Earnshaw JJ. Randomized trial of polytetrafluoroethylene patch for recurrent great saphenous varicose veins. *Eur J Vasc Endovasc Surg*. 2007;34:367-373.

66. Abela R, Liamis A, Prionidis I, et al. Reverse foam sclerotherapy of the great saphenous vein and sapheno-femoral ligation compared to standard and invagination stripping: a prospective clinical series. *Eur J Vasc Endovasc Surg*. 2008;36:485-490.

67. Winterborn RJ, Foy C, Heather H, Earnshaw JJ. Randomized trial of flush saphenofemoral ligation for primary great saphenous varicose veins. *Eur J Vasc Endovasc Surg*. 2008;36:477-484.

68. Mekako AI, Chetter IC, Coughlin PA, Hatfield J, McCollum PT; Hull Antibiotic pRophylaxis in varicose Vein Surgery Trialists (HARVEST). Randomized clinical trial of co-amoxiclav versus no antibiotic prophylaxis in varicose vein surgery. *Br J Surg*. 2010;97:29-36.

69. Haas E, Burkhardt T, Maile N. Rezidivhäufigkeit durch neoangiogenese nach modifizierter krossektomie prospektiv-randomisierte, farbduplex-kontrollierte studie. *Phlebologie*. 2005;34:101-104.

70. Carandina S, Mari C, De Palma M, et al. Varicose vein stripping vs haemodynamic correction (CHIVA): a long term randomised trial. *Eur J Vasc Endovasc Surg*. 2008;35:230-237.

71. Parés JO, Juan J, Tellez R, et al. Varicose vein surgery: stripping versus the CHIVA method—a randomized controlled trial. *Ann Surg*. 2010;251:624-631.

72. Hinchliffe RJ, Ubhi J, Beech A, Ellison J, Braithwaite BD. A prospective randomised controlled trial of VNUS closure versus surgery for the treatment of recurrent long saphenous varicose veins. *Eur J Vasc Endovasc Surg*. 2006;31:212-218.

73. Kianifard B, Holdstock JM, Whiteley MS. Radiofrequency ablation (VNUS closure) does not cause neo-vascularisation at the groin at one year: results of a case controlled study. *Surgeon*. 2006;4:71-74.

74. Lurie F, Creton D, Eklof B, et al. Prospective randomized study of endovenous radiofrequency obliteration (closure procedure) versus ligation and stripping in a selected patient population (EVOLVeS Study). *J Vasc Surg*. 2003;38:207-214.

75. Lurie F, Creton D, Eklof B, et al. Prospective randomized study of endovenous radiofrequency obliteration (closure) versus ligation and vein stripping (EVOLVeS): two-year follow-up. *Eur J Vasc Endovasc Surg*. 2005;29:67-73.

76. Rautio T, Ohinmaa A, Perälä J, et al. Endovenous obliteration versus conventional stripping operation in the treatment of primary varicose veins: a randomized controlled trial with comparison of the costs. *J Vasc Surg*. 2002;35:958-965.

77. Perälä J, Rautio T, Biancari F, et al. Radiofrequency endovenous obliteration versus stripping of the long saphenous vein in the management of primary varicose veins: 3-year outcome of a randomized study. *Ann Vasc Surg*. 2005;19:669-672.

78. Stötter L, Schaaf I, Bockelbrink A. Comparative outcomes of radiofrequency endoluminal ablation, invagination stripping and cryostripping in the treatment of great saphenous vein insufficiency. *Phlebology*. 2006;21:60-64.

79. Subromania S, Lees T. Radiofrequency ablation vs conventional surgery for varicose veins—a comparison of treatment costs in a randomized trial. *Eur J Vasc Endovasc Surg*. 2010;39:104-111.

80. Elkaffas KH, Elkashef O, ElBaz W. Great saphenous vein radiofrequency ablation versus standard stripping in the management of primary varicose veins—a randomized clinical trial. *Angiology*. 2011;62:49-54.

81. de Medeiros CA, Luccas GC. Comparison of endovenous treatment with an 810 nm laser versus conventional stripping of the great saphenous vein in patients with primary varicose veins. *Dermatol Surg*. 2005;31:1685-1694.

82. Vuylsteke M, Van den Bussche D, Audenaert EA, Lissens P. Endovenous laser obliteration for the treatment of primary varicose veins. *Phlebology*. 2006;21:80-87.

83. Lin Y, Ye CS, Huang XL, Ye JL, Yin HH, Wang SM. A random, comparative study on endovenous laser therapy and saphenous veins stripping for the treatment of great saphenous vein incompetence [in Chinese]. *Zhonghua Yi Xue Za Zhi*. 2007;87:3043-3046.

84. Rasmussen LH, Bjoern L, Lawaetz M, Blemings A, Lawaetz B, Eklof B. Randomized trial comparing endovenous laser ablation of the great saphenous vein with high ligation and stripping in patients with varicose veins: short-term results. *J Vasc Surg*. 2007;46:308-315.

85. Darwood RJ, Theivacumar N, Dellagrammaticas D, Mavor AI, Gough MJ. Randomized clinical trial comparing endovenous laser ablation with surgery for the treatment of primary great saphenous varicose veins. *Br J Surg*. 2008;95:294-301.

86. Kalteis M, Berger I, Messie-Werndl S, et al. High ligation combined with stripping and endovenous laser ablation of the great saphenous vein: early results of a randomized controlled study. *J Vasc Surg*. 2008;47:822-829.

87. Theivacumar NS, Darwood R, Gough MJ. Neovascularization and recurrence 2 years after varicose vein treatment for sapheno-femoral and great saphenous reflux: a comparison of surgery and endovenous laser ablation. *Eur J Vasc Endovasc Surg*. 2009;38:203-207.

88. Christenson JT, Gueddi S, Gemayel G, Bounameaux H. Prospective randomized trial comparing endovenous laser ablation and surgery for treatment of primary great saphenous varicose veins with a 2-year follow-up. *J Vasc Surg*. 2010;52:1234-1241.

89. Pronk P, Gauw SA, Mooij MC, et al. Randomised controlled trial comparing sapheno-femoral ligation and stripping of the great saphenous vein with endovenous laser ablation (980 nm) using local tumescent anaesthesia: one year results. *Eur J Vasc Endovasc Surg*. 2010;40:649-656.

90. Rasmussen LH, Bjoern L, Lawaetz M, Lawaetz B, Blemings A, Eklöf B. Randomized clinical trial comparing endovenous laser ablation with stripping of the great saphenous vein: clinical outcome and recurrence after 2 years. *Eur J Vasc Endovasc Surg*. 2010;39:630-635.

91. Carradice D, Mekako AI, Mazari FA, Samuel N, Hatfield J, Chetter IC. Randomized clinical trial of endovenous laser ablation compared with conventional surgery for great saphenous varicose veins. *Br J Surg*. 2011;98:501-510.

92. Carradice D, Mekako AI, Mazari FA, Samuel N, Hatfield J, Chetter IC. Clinical and technical outcomes from a randomized clinical trial of endovenous laser ablation compared with conventional surgery for great saphenous varicose veins. *Br J Sur*. 2011;98:1117-1123.

93. Rass K, Frings N, Glowacki P, et al. Comparable effectiveness of endovenous laser ablation and high ligation with stripping of the great saphenous vein: two-year results of a randomized clinical trial (RELACS study). *Arch Dermatol*. 2012;148:49-58.

94. Rasmussen L, Lawaetz M, Bjoern L, Blemings A, Eklof B. Randomized clinical trial comparing endovenous laser ablation and stripping of the great saphenous vein with clinical and duplex outcome after 5 years. *J Vasc Surg*. 2013;58:421-426.

95. Flessenkämpfer I, Hartmann M, Stenger D, Roll S. Endovenous laser ablation with and without high ligation compared with high ligation and stripping in the treatment of great saphenous varicose veins: initial results of a multicentre randomized controlled trial. *Phlebology*. 2013;28:16-23.

96. Samuel N, Carradice D, Wallace T, Mekako A, Hatfield J, Chetter I. Randomized clinical trial of endovenous laser ablation versus conventional surgery for small saphenous varicose veins. *Ann Surg*. 2013;257:419-426.

97. Roopram AD, Lind MY, Van Brussel JP, et al. Endovenous laser ablation versus conventional surgery in the treatment of small saphenous vein incompetence. *J Vasc Surg: Venous Lym Dis*. 2013;1:357-363.

98. Dzieciuchowicz L, Espinosa G, Páramo JA. Hemostatic activation and inflammatory response after three methods of treatment of great saphenous vein incompetence. *Phlebology*. 2014;29:154-163.

99. Einarsson E, Eklöf B, Neglén P. Sclerotherapy or surgery as treatment for varicose veins: a prospective randomized study. *Phlebology*. 1993;8:22-26.

100. Rutgers PH, Kitslaar PJ. Randomized trial of stripping versus high ligation combined with sclerotherapy in the treatment of the incompetent greater saphenous vein. *Am J Surg*. 1994;168:311-315.

101. Belcaro G, Nicolaides AN, Ricci A, et al. Endovascular sclerotherapy, surgery, and surgery plus sclerotherapy in superficial venous incompetence: a randomized, 10-year follow-up trial—final results. *Angiology*. 2000;51:529-534.

102. Belcaro G, Cesarone MR, Di Renzo A, et al. Foam-sclerotherapy, surgery, sclerotherapy, and combined treatment for varicose veins: a 10-year, prospective, randomized, controlled, trial (VEDICO trial). *Angiology*. 2003;54:307-315.

103. de Roos KP, Nieman FH, Neumann HA. Ambulatory phlebectomy versus compression sclerotherapy: results of a randomized controlled trial. *Dermatol Surg*. 2003;29:221-226.

104. Bountouroglou DG, Azzam M, Kakkos SK, Pathmarajh M, Young P, Geroulakos G. Ultrasound-guided foam sclerotherapy combined with sapheno-femoral ligation compared to surgical treatment of varicose veins: early results of a randomised contolled trial. *Eur J Vasc Endovasc Surg*. 2006;31:93-100.

105. Liu X, Jia X, Guo W, et al. Ultrasound-guided sclerotherapy of the great saphenous vein with sapheno-femoral ligation compared to standard stripping: a prospective clinical study. *Int Angiol*. 2011;30:321-326.

106. Kalodiki E, Lattimer CR, Azzam M, Shawish E, Bountouroglou D, Geroulakos G. Long-term results of a randomized controlled trial on ultrasound-guided foam sclerotherapy combined with saphenofemoral ligation vs standard surgery for varicose veins. *J Vasc Surg*. 2012;55:451-457.

107. Figueiredo M, Araújo S, Barros N Jr, Miranda F Jr. Results of surgical treatment compared with ultrasound-guided foam sclerotherapy in patients with varicose veins: a prospective randomised study. *Eur J Vasc Endovasc Surg*. 2009;38:758-763.

108. Shadid N, Ceulen R, Nelemans P, et al. Randomized clinical trial of ultrasound-guided foam sclerotherapy versus surgery for the incompetent great saphenous vein. *Br J Surg.* 2012;99:1062-1070.

109. Wright D, Gobin JP, Bradbury AW, et al; Varisolve® European Phase III Investigators Group. Varisolve® polidocanol microfoam compared with surgery or sclerotherapy in the management of varicose veins in the presence of trunk vein incompetence: European randomized controlled trial. *Phlebology*. 2006;21:180-190.

110. Biemans AA, Kockaert M, Akkersdijk GP, et al. Comparing endovenous laser ablation, foam sclerotherapy, and conventional surgery for great saphenous varicose veins. *J Vasc Surg*. 2013;58:727-734.

111. Brittenden J, Cotton SC, Elders A, et al. A randomized trial comparing treatments for varicose veins. *N Engl J Med*. 2014;371:1218-1227.

112. Tassie E, Scotland G, Brittenden J, et al; CLASS Study Team. Cost-effectiveness of ultrasound-guided foam sclerotherapy, endovenous laser ablation or surgery as treatment for primary varicose veins from the randomized CLASS trial. *Br J Surg*. 2014;101:1532-1540.

113. Rasmussen LH, Lawaetz M, Bjoern L, Vennits B, Blemings A, Eklof B. Randomized clinical trial comparing endovenous laser ablation, radiofrequency ablation, foam sclerotherapy and surgical stripping for great saphenous varicose veins. *Br J Surg*. 2011;98:1079-1087.

114. Rasmussen L, Lawaetz M, Serup J, et al. Randomized clinical trial comparing endovenous laser ablation, radiofrequency ablation, foam sclerotherapy and surgical stripping for great saphenous varicose veins with 3-year follow-up. *J Vasc Surg: Venous Lym Dis*. 2013;1:349-356.

115. Woźniak W, Mlosek RK, Ciostek P. Assessment of the efficacy and safety of steam vein sclerosis as compared to classic surgery in lower extremity varicose vein management. *Wideochir Inne Tech Maloinwazyjne*. 2015;10:15-24.

116. Campanello M, Hammarsten J, Forsberg C, Bernland P, Henrikson O, Jensen J. Standard stripping versus long saphenous vein–saving surgery for primary varicose veins: a prospective, randomized study with the patients as their own controls. *Phlebology*. 1996;11:45-49.

117. Hammarsten J, Pederson P, Cederlund CG, Campanello M. Long saphenous vein saving surgery for varicose veins: a long-term follow-up. *Eur J Vasc Surg*. 1990;4:361-364.

118. Hammarsten J, Campanello M, Pederson P. Long saphenous vein saving surgery for varicose veins. *Eur J Vasc Surg*. 1993;7:763-764.

119. Winterborn RJ, Foy C, Earnshaw JJ. Causes of varicose vein recurrence: late results of a randomized controlled trial of stripping the long saphenous vein. *J Vasc Surg*. 2004;40:634-639.

120. Almeida JI, Kaufman J, Göckeritz O, et al. Radiofrequency endovenous ClosureFAST versus laser ablation for the treatment of great saphenous reflux: a multicenter, single-blinded, randomized study (RECOVERY study). *J Vasc Interv Radiol*. 2009;20:752-759.

121. Shepherd AC, Gohel MS, Brown LC, Metcalf MJ, Hamish M, Davies AH. Randomized clinical trial of VNUS ClosureFAST radiofrequency ablation versus laser for varicose veins. *Br J Surg*. 2010;97:810-818.

122. Gale SS, Lee JN, Walsh ME, Wojnarowski DL, Comerota AJ. A randomized, controlled trial of endovenous thermal ablation using the 810-nm wavelength laser and the ClosurePLUS radiofrequency ablation methods for superficial venous insufficiency of the great saphenous vein. *J Vasc Surg*. 2010;52:645-650.

123. Goode SD, Chowdhury A, Crockett M, et al. Laser and radiofrequency ablation study (LARA study): a randomised study comparing radiofrequency ablation and endovenous laser ablation (810 nm). *Eur J Vasc Endovasc Surg*. 2010;40:246-253.

124. Nordon IM, Hinchliffe RJ, Brar R, et al. A prospective double-blind randomized controlled trial of radiofrequency versus laser treatment of the great saphenous vein in patients with varicose veins. *Ann Surg*. 2011;254:876-881.

125. Doganci S, Demirkilic U. Comparison of 980 nm laser and bare-tip fibre with1470 nm laser and radial fibre in the treatment of great saphenous vein varicosities: a prospective randomized controlled trial. *Eur J Vasc Endovasc Surg*. 2010;40:254-259.

126. Lane TR, Kelleher D, Shepherd AC, Franklin IJ, Davies AH. Ambulatory varicosity avulsion later or synchronized (AVULS): a randomized clinical trial. *Ann Surg*. 2015;261:654-661.

127. Disselhoff BC, der Kinderen DJ, Moll FL. Is there a risk for lymphatic complications after endovenous laser treatment versus cryostripping of the great saphenous vein? A prospective study. *Phlebology*. 2008;23:10-14.

128. Disselhoff BC, der Kinderen DJ, Kelder JC, Moll FL. Randomized clinical trial comparing endovenous laser with cryostripping for great saphenous varicose veins. *Br J Surg*. 2008;95:1232-1238.

129. Disselhoff BC, Buskens E, Kelder JC, der Kinderen DJ, Moll FL. Randomized comparison of costs and cost-effectiveness of cryostripping and endovenous laser ablation for varicose veins: 2-year results. *Eur J Vasc Endovasc Surg*. 2009;37:357-363.

130. Lattimer CR, Kalodiki E, Azzam M, Geroulakos G. Validation of a new duplex derived haemodynamic effectiveness score, the saphenous treatment score, in quantifying varicose vein treatments. *Eur J Vasc Endovasc Surg*. 2012;43:348-354.

131. Lattimer CR, Azzam M, Kalodiki E, Shawish E, Trueman P, Geroulakos G. Cost and effectiveness of laser with phlebectomies compared with foam sclerotherapy in superficial venous insufficiency. Early results of a randomised controlled trial. *Eur J Vasc Endovasc Surg*. 2012;43:594-600.

132. Lattimer CR, Kalodiki E, Azzam M, Makris GC, Somiayajulu S, Geroulakos G. Interim results on abolishing reflux alongside a randomized clinical trial on laser ablation with phlebectomies versus foam sclerotherapy. *Int Angiol*. 2013;32:394-403.

133. Kabnick LS. Outcome of different endovenous laser wavelengths for great saphenous vein ablation. *J Vasc Surg*. 2006;43:88-93.

134. Disselhoff BC, der Kinderen DJ, Kelder JC, Moll FL. Randomized clinical trial comparing endovenous laser ablation of the great saphenous vein with and without ligation of the sapheno-femoral junction: 2-year results. *Eur J Vasc Endovasc Surg.* 2008;36:713-718.

135. Disselhoff BC, der Kinderen DJ, Kelder JC, Moll FL. Five-year results of a randomised clinical trial of endovenous laser ablation of the great saphenous vein with and without ligation of the saphenofemoral junction. *Eur J Vasc Endovasc Surg.* 2011;41:685-690.

136. Theivacumar NS, Dellagrammaticas D, Mavor AI, Gough MJ. Endovenous laser ablation: does standard above-knee great saphenous vein ablation provide optimum results in patients with above- and below-knee reflux? A randomized controlled trial. *J Vasc Surg.* 2008;48:173-178.

137. Hogue RS, Schul MW, Dando CF, Erdman BE. The effect of nitroglycerin ointment on great saphenous vein targeted venous access size diameter with endovenous laser treatment. *Phlebology.* 2008;23:222-226.

138. Pannier F, Rabe E, Maurins U. 1470 nm diode laser for endovenous ablation (endovenous laser ablation) of incompetent saphenous veins—a prospective randomized pilot study comparing warm and cold tumescence anesthesia. *Vasa.* 2010;39:249-255.

139. Dumantepe M, Uyar I. Comparing cold and warm tumescent anesthesia for pain perception during and after the endovenous laser ablation procedure with 1470 nm diode laser. *Phlebology.* 2015;30:45-51.

140. Vuylsteke M, De Bo T, Dompe G, Di Crisci D, Abbad C, Mordon S. Endovenous laser treatment: is there a clinical difference between using a 1500 nm and a 980 nm diode laser? A multicenter randomised clinical trial. *Int Angiol.* 2011;30:327-334.

141. Bakker NA, Schieven LW, Bruins RM, van den Berg M, Hissink RJ. Compression stockings after endovenous laser ablation of the great saphenous vein: a prospective randomized controlled trial. *Eur J Vasc Endovasc Surg.* 2013;46:588-592.

142. Samuel N, Wallace T, Carradice D, Mazari FA, Chetter IC. Comparison of 12-w versus 14-w endovenous laser ablation in the treatment of great saphenous varicose veins: 5-year outcomes from a randomized controlled trial. *Vasc Endovascular Surg.* 2013;47:346-352.

143. Kahle B, Leng K. Efficacy of sclerotherapy in varicose veins—a prospective, blinded, placebo-controlled study. *Dermatol Surg.* 2004;30:723-728.

144. Hamel-Desnos C, Desnos P, Wollmann JC, Ouvry P, Mako S, Allaert FA. Evaluation of the efficacy of polidocanol in the form of foam compared with liquid form in sclerotherapy of the greater saphenous vein: initial results. *Dermatol Surg.* 2003;29:1170-1175.

145. Yamaki T, Nozaki M, Iwasaki S. Comparative study of duplex-guided foam sclerotherapy and duplex-guided liquid sclerotherapy for the treatment of superficial venous insufficiency. *Dermatol Surg.* 2004;30:718-722.

146. Alòs J, Carreño P, López JA, Estadella B, Serra-Prat M, Marinel-Lo J. Efficacy and safety of sclerotherapy using polidocanol foam: a controlled clinical trial. *Eur J Vasc Endovasc Surg.* 2006;31:101-107.

147. Ouvry P, Allaert FA, Desnos P, Hamel-Desnos C. Efficacy of polidocanol foam versus liquid in sclerotherapy of the great saphenous vein: a multicentre randomised controlled trial with a 2-year follow-up. *Eur J Vasc Endovasc Surg.* 2008;36:366-370.

148. Rabe E, Otto J, Schliephake D, Pannier F. Efficacy and safety of great saphenous vein sclerotherapy using standardised polidocanol foam (ESAF): a randomised controlled multicentre clinical trial. *Eur J Vasc Endovasc Surg.* 2008;35:238-245.

149. Hamel-Desnos C, Allaert FA, Benigni JP, et al; Société Française de Phlébologie. Study 3/1. Polidocanol foam 3% versus 1% in the great saphenous vein: early results [in French]. *Phlébologie.* 2005;58:165-173.

150. Ceulen RP, Bullens-Goessens YI, Pi-Van De Venne SJ, Nelemans PJ, Veraart JC, Sommer A. Outcomes and side effects of duplex-guided sclerotherapy in the treatment of great saphenous veins with 1% versus 3% polidocanol foam: results of a randomized controlled trial with1-year follow-up. *Dermatol Surg.* 2007;33:276-281.

151. Hamel-Desnos C, Ouvry P, Benigni JP, et al. Comparison of 1% and 3% polidocanol foam in ultrasound guided sclerotherapy of the great saphenous vein: a randomised, double-blind trial with 2 year-follow-up: "the 3/1 study." *Eur J Vasc Endovasc Surg*. 2007;34:723-729.

152. Blaise S, Bosson JL, Diamand JM. Ultrasound-guided sclerotherapy of the great saphenous vein with 1% vs. 3% polidocanol foam: a multicentre double-blind randomised trial with 3-year follow-up. *Eur J Vasc Endovasc Surg*. 2010;39:779-786.

153. O'Hare JL, Stephens J, Parkin D, Earnshaw JJ. Randomized clinical trial of different bandage regimens after foam sclerotherapy for varicose veins. *Br J Surg*. 2010;97:650-656.

154. Hamel-Desnos CM, Guias BJ, Desnos PR, Mesgard A. Foam sclerotherapy of the saphenous veins: randomized controlled trial with or without compression. *Eur J Vasc Endovasc Surg*. 2010;39:500-507.

155. Hamel-Desnos CM, Desnos PR, Ferre B, Le Querrec A. In vivo biological effects of foam sclerotherapy. *Eur J Vasc Endovasc Surg*. 2011;42:238-245.

156. Yamaki T, Hamahata A, Soejima K, Kono T, Nozaki M, Sakurai H. Prospective randomised comparative study of visual foam sclerotherapy alone or in combination with ultrasound-guided foam sclerotherapy for treatment of superficial venous insufficiency: preliminary report. *Eur J Vasc Endovasc Surg*. 2012;43:343-347.

157. Hamel-Desnos CM, Gillet JL, Desnos PR, Allaert FA. Sclerotherapy of varicose veins in patients with documented thrombophilia: a prospective controlled randomized study of 105 cases. *Phlebology*. 2009;24:176-182.

158. Eklof B, Perrin M, Delis K, Rutherford RB, Gloviczki P. Updated terminology of chronic venous disorders: the VEIN-TERM transatlantic interdisciplinary consensus document. *J Vasc Surg*. 2009;49:498-501.

159. Perrin MR, Guex JJ, Ruckley CV, et al; REVAS Group. Recurrent varices after surgery (REVAS), a consensus document. *Cardiovasc Surg*. 2000;8:233-245.

160. Vasquez MA, Rabe E, McLafferty RB, et al; American Venous Forum Ad Hoc Outcomes Working Group. Revision of the venous clinical severity score: venous outcomes consensus statement. *J Vasc Surg*. 2010;52:1387-1396.

161. Vasquez MA, Munschauer CE. Venous clinical severity score and quality-of-life assessment tools: application to vein practice. *Phlebology*. 2008;23:259-275.

162. Shepherd AC, Gohel MS, Lim CS, Davies AH. A study to compare disease-specific quality of life with clinical anatomical and hemodynamic assessments in patients with varicose veins. *J Vasc Surg*. 2011;53:374-382.

163. Guex JJ. Patient-reported outcome or physician-reported outcome? *Phlebology*. 2008;23:251.

164. Fischer R, Linde N, Duff C. Cure and reappearance of symptoms of varicose veins after stripping operation: a 34 year follow-up. *J Phlebology*. 2001;1:49-60.

165. van Rij AM, Jiang P, Solomon C, Christie RA, Hill GB. Recurrence after varicose vein surgery: a prospective long-term clinical study with duplex ultrasound scanning and air plethysmography. *J Vasc Surg*. 2003;38:935-943.

166. Kostas T, Ioannou CV, Toulouopakis E, et al Recurrent varicose veins after surgery: a new appraisal of a common and complex problem in vascular surgery. *Eur J Vasc Endovasc Surg*. 2004;27:275-282.

167. Merchant RF, Pichot O; Closure Study Group. Long-term outcomes of endovenous radiofrequency obliteration of saphenous reflux as a treatment for superficial venous insufficiency. *J Vasc Surg*. 2005;42:502-509.

168. van den Bos R, Arends L, Kockaert M, Neumann M, Nijsten T. Endovenous therapies of lower extremity varicosities: a meta-analysis. *J Vasc Surg*. 2009;49:230-239.

169. Brar R, Nordon IM, Hinchliffe RJ, Loftus IM, Thompson MM. Surgical management of varicose veins: meta-analysis. *Vascular*. 2010;18:205-220.

170. Murad MH, Coto-Yglesias F, Zumaeta-Garcia M, et al. A systematic review and meta-analysis of the treatments of varicose veins. *J Vasc Surg*. 2011;53(suppl 5):49S-65S.

171. Nesbitt C, Eifell RK, Coyne P, Badri H, Bhattacharya V, Stansby G. Endovenous ablation (radiofrequency and laser) and foam sclerotherapy versus conventional surgery for great saphenous vein varices. *Cochrane Database Syst Rev.* 2011:CD005624.

172. Tellings SS, Ceulen RP, Sommer A. Surgery and endovenous techniques for the treatment of small saphenous varicose veins: a review of the literature. *Phlebology.* 2011;26:179-184.

173. Siribumrungwong B, Noorit P, Wilasrusmee C, Attia J, Thakkinstian A. A systematic review and meta-analysis of randomised controlled trials comparing endovenous ablation and surgical intervention in patients with varicose vein. *Eur J Vasc Endovasc Surg.* 2012;44:214-223.

174. Puggioni A, Lurie F, Kistner RL, Eklof B. How often is deep venous reflux eliminated after saphenous vein ablation? *J Vasc Surg.* 2003;38:517-521.

175. Guarnera G, Furgiuele S, Di Paola FM, Camilli S. Recurrent varicose veins and primary deep venous insufficiency: relationship and therapeutic implications. *Phlebology.* 1995;10:98-102.

176. Perrin MR. Results of deep-vein reconstruction. *Vasc Endovasc Surg.* 1997;31:273-275.

177. Campbell WA, West A. Duplex ultrasound audit of operative treatment of primary varicose veins. In: Negus D Jantet G, Coleridge-Smith PD, eds. *Phlebology '95.* London, UK: Springer; 1995:407-409.

178. Stuart WP, Adam DJ, Allan PL, Ruckley CV, Bradbury AW. Saphenous surgery does not correct perforator incompetence in the presence of deep venous reflux. *J Vasc Surg.* 1998;28:834-838.

179. Al-Mulhim AS, El-Hoseiny H, Al-Mulhim FM, et al. Surgical correction of main stem reflux in the superficial venous system: does it improve the blood flow of incompetent perforating veins? *World J Surg.* 2003;27:793-796.

180. Mendes RR, Marston WA, Farber MA, Keagy BA. Treatment of superficial and perforator venous incompetence without deep venous insufficiency: is routine perforator ligation necessary? *J Vasc Surg.* 2003;38:891-895.

181. Perrin M. Presence of varices after operative treatment: a review (Part 2). *Phlebolymphology.* 2015;22:5-11.

182. van Gent WB, Wittens CHA. Influence of perforating vein surgery in patients with venous ulceration. *Phlebology.* 2015;30:127-132.

183. Kakkos SK, Bountouroglou DG, Azzam M, Kalodiki E, Daskalopoulos M, Geroulakos G. Effectiveness and safety of ultrasound-guided foam sclerotherapy for recurrent varicose veins: immediate results. *J Endovasc Ther.* 2006;13:357-364.

184. Darvall KA, Bate GR, Adam DJ, Silverman SH, Bradbury AW. Duplex ultrasound outcomes following ultrasound-guided foam sclerotherapy of symptomatic recurrent great saphenous varicose veins. *Eur J Vasc Endovasc Surg.* 2011;42:107-114.

185. Gloviczki P, Comerota AJ, Dalsing MC, et al. The care of patients with varicose veins and associated chronic venous diseases: clinical practice guidelines of the Society for Vascular Surgery and the American Venous Forum. *J Vasc Surg.* 2011;53(suppl 5):2S-48S.

186. Lugli M, Maleti O, Perrin M. Review and comment of the 2011 clinical practice guidelines of the Society for Vascular Surgery and the American Venous Forum. *Phlebolymphology.* 2012;19:107-120.

187. Nicolaides A, Kakkos S, Eklof B, et al. Management of chronic venous disorders of the lower limbs—guidelines according to scientific evidence. *Int Angiol.* 2014;33:87-208.

188. Pavlović MD, Petrović SS, Pichot O, et al. Guidelines of the First International Consensus Conference on Endovenous Thermal Ablation for Varicose Vein Disease: ETAV Consensus Meeting 2012. *Phlebology.* 2015;30:257-273.

189. Wittens C, Davies AH, Baekgaard N, et al. Management of chronic venous disease: clinical practice guidelines of the European Society for Vascular Surgery (ESVS). *Eur J Vasc Endovasc Surg.* 2015;49:678-737.

190. Guyatt G, Gutterman D, Baumann MH, et al. Grading strength of recommendations and quality of evidence in clinical guidelines: report from an American College of Chest Physicians Task Force. *Chest.* 2006;129:174-181.

191. Perk J, De Backer G, Gohlke H, et al. European guidelines on cardiovascular disease prevention in clinical practice (version 2012). *Eur Heart J.* 2012;33:1635-1701.

192. National Institute for Health and Care Excellence (NICE). Varicose veins: diagnosis and management. NICE guidelines [CG168]. Available at: http://www.nice.org.uk/guidance/cg168/chapter/1-recommendations. Published July 2013. Accessed January 19, 2016.

193. National Institute for Health and Care Excellence (NICE). Radiofrequency ablation of varicose veins. NICE interventional procedure guidance 8 [ipg8]. Available at: http://www.nice.org.uk/guidance/ipg8. Published September 2003. Accessed January 19, 2016.

194. National Institute for Health and Care Excellence (NICE). Endovenous laser treatment of the long saphenous vein. NICE interventional procedure guidance 52 [ipg52]. Available at: http://www.nice.org.uk/guidance/ipg52. Published March 2004. Accessed January 19, 2016.

195. National Institute for Health and Care Excellence (NICE). Ultrasound-guided foam sclerotherapy for varicose veins. NICE interventional procedure guidance 440 [ipg440]. Available at: http://www.nice.org.uk/guidance/ipg440. Published February 2013. Accessed January 19, 2016.