Retrograde Transpedal Access for Revascularization of Belowthe-Knee Arteries in Patients with Critical Limb Ischemia after an Unsuccessful Antegrade Transfemoral Approach

Retrograd transpedale Revaskularisierung kruraler Arterien bei Patienten mit kritischer Extremitätenischämie und gescheitertem antegraden Revaskularisierungsversuch

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Key words

- CLI
- pedal access
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- retrograde
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Zusammenfassung



Ziel: Evaluation der Sicherheit sowie technischem und klinischem Erfolg der endovaskulären Revaskularisierung kruraler Gefäßokklusionen über einen retrograd transpedalen Zugang.

Material und Methoden: Retrospektiv wurden 172 Patienten mit einer Unterschenkelrevaskularisierung zwischen 2/2014 und 8/2015 identifiziert. Bei 16/172 Patienten (9,3%) war ein transfemoraler Rekanalisierungsversuch gescheitert und ein retrograd transpedaler Zugang unternommen worden. Unter sonografischer Führung wurden die A. dorsalis pedis (n = 13) oder die A. tibialis posterior unter Verwendung eines speziellen Sets punktiert. Die Prozedur wurde durch eine antegrade PTA beendet. Analysiert wurden Komorbiditäten sowie der Gefäßdurchmesser und -kalzifizierungsgrad des punktierten Gefäßes. Endpunkte waren technischer und prozeduraler Erfolg, periprozedurale Komplikationen, Prozedurzeit und Extremitätenerhalt.

Ergebnisse: Bei 15 Patienten (93,8%) fanden sich jeweils Diabetes, KHK und Hypertonie unter den Komorbiditäten. Nierenfunktionseinschränkungen und stattgehabte Amputation fanden sich bei 7 Patienten (43,8%). Bei 5/16 Patienten (31,3%) bestand eine Kalzifizierung des punktierten Gefäßen, der mittlere Durchmesser betrug 1,75 +/-0,24 mm. Die Prozedurdauer betrug 92,4+/-23 min. Der technische Erfolg betrug 100%. Die retrograde Läsionspassage war bei 12/16 Patienten (75,0%), die Gesamtprozedur bei 10/16 Patienten (68,8%) erfolgreich. Minorkomplikationen kamen bei 2/16 Patienten (12,5%) vor. Der Extremitätenerhalt, bzw. Überlebensrate nach 12 Monaten betrugen 72,9% und 100%. Majoramputationen nach der Revaskularisierung wurden bei 2/16 Patienten (12,5%) durchgeführt.

Schlussfolgerung: Nach gescheitertem antegraden Revaskularisierungsversuch kann ein retrograd transpedaler Zugang den Prozedurerfolg er-

Abstract



Purpose: To evaluate the safety and technical and clinical success of endovascular below-the-knee (BTK) artery revascularization by a retrograde transpedal access.

Materials and Methods: We retrospectively identified 16/172 patients (9.3%) with endovascular BTK revascularization in whom a transfemoral approach had failed and transpedal access had been attempted. The dorsal pedal (n=13) or posterior tibial (n=3) artery was accessed using a dedicated access set and ultrasound guidance. The procedure was finished in antegrade fashion by plain old balloon angioplasty (POBA). Comorbidities, vessel diameter and calcification at the access site were recorded. The analyzed outcomes were technical success, procedural complications, procedure time, crossing (guidewire beyond lesion and intra-luminal) and procedural (residual stenosis <30% after POBA) success, and limb salvage.

Results: Diabetes, coronary artery disease and hypertension were present in 15 patients (93.8%), and both renal impairment and previous amputations in 7 (43.8%). Pedal access vessel calcification was present in 5/16 patients (31.3%). The mean diameter was 1.75+/-0.24 mm. The procedure time was 92.4+/-23 min. The success rate for achieving retrograde access was 100%. Retrograde crossing was successful in 12/16 patients (75.0%). Procedural success was observed in 10/16 patients (68.8%). Minor complications occurred in 2/16 patients (12.5%). The rate of limb salvage was 72.9%, and the overall survival was 100% at 12 months. Major amputations after revascularization occurred in 2/16 patients (12.5%).

Conclusion: If an antegrade transfemoral approach to BTK lesions fails, a retrograde transpedal approach may nevertheless facilitate treatment. This approach appears to be safe and offers high technical and acceptable clinical success rates.

möglichen. Die retrograde Punktion erscheint sicher und mit hoher technischer und akzeptabler klinischer Erfolgsrate vergesellschaftet zu sein.

Kernaussagen:

- ► Retrograd transpedale Gefäßzugänge erscheinen sicher und mit hoher technischer Erfolgsrate nutzbar zu sein
- ▶ Nach technisch erfolgreicher retrograder Punktion kann bei subintimaler Läsionpassage das Reentry problematisch sein
- Nach gescheitertem antegraden Revaskularisierungsversuch verspricht die retrograde transpedale Punktion Erfolg

Key Points:

- Retrograde approaches via transpedal or transtibial vessels are safe and offer high technical success.
- ▶ One problem after technically successful puncture might be the re-entry following subintimal retrograde lesion crossing.
- ➤ After a failed attempt at antegrade revascularization of a BTK occlusion, a retrograde approach should be performed.

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Introduction



In addition to other supporting therapies, patients suffering from critical limb ischemia (CLI) with tissue loss require immediate revascularization in order to stimulate ulcer healing and minimize the risk of (major) amputation [1, 2]. Venous (popliteal-pedal) bypass has been advocated as the revascularization method of choice for below-the-knee (BTK) lesions owing to the acceptable patency rates [3]. However, in CLI patients limb salvage has been advocated as the major goal of revascularization with patency rates being less important [4], and many experienced centers prefer an endovascular approach first [5]. Reasons for this include but may not be limited to the minimally invasive character of the endovascular approach. The advantages of the endovascular approach include lower complication rates and perioperative risk due to comorbidities, no issues with a lack of conduits and cost-effectiveness compared to bypass surgery [1, 6].

As with all vascular lesions, a prerequisite of endovascular treatment is lesion crossing with the guidewire tip in an intraluminal position beyond it to start the revascularization procedure [7]. Crossing those lesions may be difficult, last but not least owing to the calcification of BTK arteries which is observed with above average frequency in diabetic and end-stage renal disease patients [5]. Furthermore, as reported about the coronaries, the proximal cap of a lesion might be more solid than the distal one owing to the fibrous tissue content [8]. This might explain why even using modern equipment – approximately 20% of tibial lesions may not be tackled successfully from an antegrade transfemoral route [9]. In this context retrograde transpedal and transtibial access has been described to overcome this problem and initial data have shown promising results regarding technical success, safety and limb salvage [1, 5, 10, 11].

As the literature regarding retrograde transpedal access is limited, we found it of interest to report our initial retrospective results with this technique using ultrasound guidance and a dedicated pedal access set for the treatment of BTK occlusions after an antegrade transfemoral approach had ultimately failed.

Materials and Methods



Study sample

We retrospectively searched through our electronic database to identify patients who had a BTK intervention in our interventional radiology department between 2/2014 and 8/2015. We identified 172 patients who underwent BTK vessel revascularization for CLI. 16/172 patients (9.3%) had a retrograde transpedal/transtibial approach after an antegrade transfemoral approach

had ultimately failed. The retrograde approach was carried out in the same session as the antegrade in all but two patients who were not able to lie in a supine position owing to discomfort. These patients had a staged procedure on days 3 and 5 after the antegrade approach had failed. All patients were discussed in our weekly interdisciplinary vascular board (interventional radiology, vascular surgery) and found to be poor candidates for either a pedal or tibial bypass. Patient and lesion characteristics are included in • Table 1.

Revascularization technique

With the patient in a supine position, we regularly perform antegrade transfemoral puncture of the common femoral artery using either fluoroscopy and/or ultrasound guidance. Using Seldinger's technique, a 10 cm long 4 – 6F introducer sheath (Radifo-

 Table 1
 Demographic data and angiographic characteristics.

age	77.7 +/-8.8y (range: 52 - 95y)
sex	10 men (62.5 %), 6 women
	(37.5%)
comorbidities	
hypertension	16/16 (100%)
 coronary artery disease 	15/16 (93.8 %)
– hyperlipidemia	15/16 (93.8 %)
 diabetes mellitus 	15/16 (93.8 %)
 chronic renal failure 	8/16 (50.0 %)
 dialysis dependency 	3/16 (18.8%)
Rutherford-Becker classification	
- category 5	10/16 (62.5 %)
- category 6	6/16 (37.5 %)
prevent III score [21]	
medium risk (4 – 7 points)	12/16 (75.0 %)
high risk (> = 8 points)	4/16 (25.0 %)
side and vessel used for retrograde a	ccess
– left	8/16 (50.0 %)
- right	8/16 (50.0 %)
 dorsal pedal artery 	13/16 (81.25%)
 distal posterior tibial artery 	3/16 (18.75 %)
lesion characteristics	
- total occlusion	16/16 (100%)
– lesion length	19.1 +/- 7.0 cm
 calcification present 	7/16 (43.8%)
Initial number of non-occluded run-	off vessels
- 0	7/16 (43.8 %)
- 1	8/16 (50.0 %)
- 2	1/16 (6.3 %)
n = 16	

n = 16.

cus®, Introducer II, Terumo, Tokyo, Japan) is inserted. 5000 units of heparin are administered via the femoral sheath. An angiogram using manual contrast injection is performed down to the foot (Fig. 1). In the case of femoral or popliteal lesions, these are addressed first, preferably by intraluminal lesion crossing. Following revascularization by plain old balloon angioplasty (POBA) in combination with either drug-eluting balloon (DEB) PTA or implantation of a stent or stentgraft, BTK lesions are addressed. As standard practice, we try to revascularize as many BTK vessels as possible in CLI patients. A 4F catheter was used to selectively intubate the diseased crural artery. In the case of an occluded vessel, we preferably use a 0.018" (V-18™ Control Wire, Boston Scientific, Marlborough, MA, USA) or 0.014" (Command ES™, Abbot Vascular, Diegem, Belgium) guidewire, supported by a vessel size-matching balloon catheter if needed. In the case of a stenosed vessel or for subintimal revascularization of an occluded one, an angled 0.018" guidewire (Radifocus®, Glidewire Advantage™, Terumo, Europe, N.V. Leuven, Belgium) is used. We do not regularly use support catheters. Following passage of the lesion, we perform POBA for 60s using either a

0.014" PTA balloon (Coyote™, Boston Scientific, Marlborough, NA, USA) or a 0.018" balloon (Saber™, Cordis, Tipperary, Ireland). In the case of failure to pass the lesion with a guidewire in an antegrade direction from above, we switch to the transpedal/ transtibial approach (o Fig. 2). Before the foot and distal part of the lower extremity are prepared by sterile washing, the initial run-off angiogram is thoroughly examined to confirm a distal vessel segment that is long enough to be punctured and in which the pedal access sheath can securely be placed. Using a linear ultrasound probe (Sparq™, Philips Healthcare) in B-mode and Duplex mode, either the dorsal pedal or distal tibial artery is identified. Using a dedicated vascular access set (Micropuncture® Pedal Introducer Access Set, Cook Medical, Bloomington, USA), the target vessel is punctured under sonographic guidance until blood flow through the 21G needle is registered. In the case of an anterior tibial artery occlusion, the dorsal pedal artery is punctured as we find it easier to access compared with the distal anterior tibial artery - probably owing to a more superficial location and fixed position. In the case of a posterior tibial artery occlusion, the distal segment near the ankle is accessed. The 0.018" set guidewire

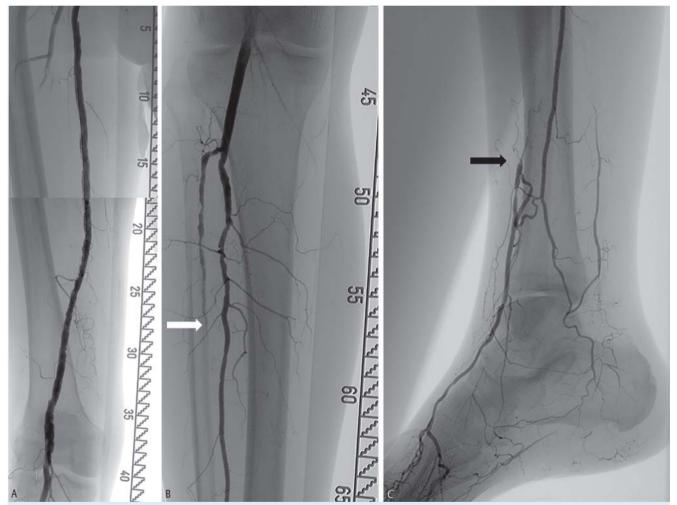


Fig. 1 Initial angiogram of a patient from our study sample with a nonhealing ulceration in a pretibial location and the dorsum of the foot. **A** DSA shows no relevant lesion of the SFA or popliteal artery in the target limb. **B, C** demonstrate one vessel run-off via the peroneal artery and chronic total occlusions of the posterior and anterior (white arrow) tibial arteries. The distal anterior tibial artery is reconstituted (black arrow) via a peroneal collateral and fills the dorsal pedal artery.

Abb. 1 Initiales Angiogramm eines Patienten mit nicht heilenden Ulzerationen prätibial und auf dem Fußrücken **A** die DSA zeigt die frei durchgängigen A. fem. sup. und A. poplitea. **B, C** zeigen eine 1-Gefäßversorgung über die A. fibularis sowie chronische Gefäßverschlüsse von A. tib. anterior (weißer Pfeil) und posterior. Rekonstitution der distalen A. tib. anterior (schwarzer Pfeil) über eine Kollaterale der A. fibularis und Kontrastierung der A. dorsalis pedis.

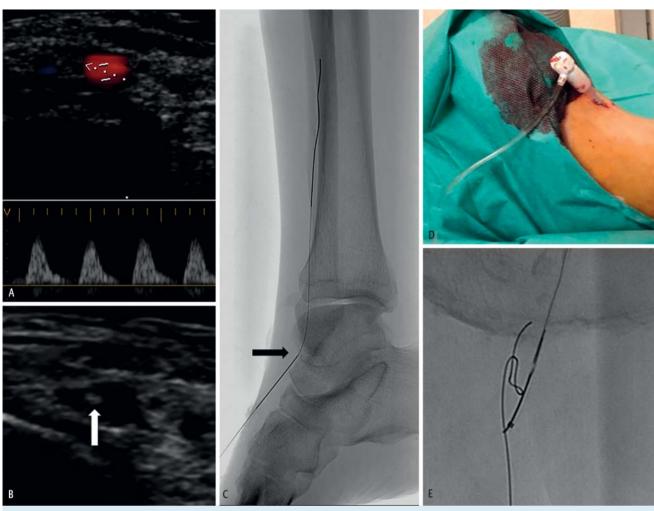


Fig. 2 Retrograde transpedal access in the patient shown in Fig. 1. **A** Duplex ultrasound is used to detect the dorsal pedal artery. **B** Using a linear probe in an axial fashion, the dorsal pedal artery is punctured with a 21G needle (white arrow). **C** A 0.018" guidewire from the pedal access introducer set is inserted via the needle (black arrow) and **D** the (4F) pedal access sheath is placed. **E** After exchanging the set wire for a CTO wire, the latter is snared in the proximal SFA and guided out via the femoral sheath.

Abb. 2 Retrograder transpedaler Gefäßzugang bei dem in Abb. 1 gezeigten Patienten. **A** Farbkodierte Dopplersonografie zur Darstellung der A. dorsalis pedis. **B** Unter Zuhilfenahme eines Linearschallkopfes (axiale Ausrichtung) wird die A. dorsalis pedis mit einer 21G starken Nadel punktiert (weißer Pfeil). **C** Der im Punktionsset enthaltene 0.018" Draht wird über die Punktionsnadel eingebracht (schwarzer Pfeil) und **D** die (4F) Schleuse aus dem pedal access set® platziert. **E** Nach Einwechseln eines CTO-Drahtes wird dieser in der proximalen A. fem. superf. mit einem Snare-Device gefasst und transfemoral ausgeleitet.

is then introduced using fluoroscopy control. Following local anesthesia, a small incision over the puncture side, and removal of the needle, the 4F set sheath is placed over the wire. This sheath comes with a (Check-Flo®) hemostasis valve. A retrograde manual angiogram via this sheath is performed to confirm the intraluminal position of the sheath as well as the configuration of the distal cap of the occlusion. We do not regularly use any intra-arterial medication unless spasm occurs. The occlusion is then traversed in a retrograde and preferably intraluminal manner using either a 0.014" or 0.018" straight CTO guidewire as mentioned above. In the case of a subintimal passage, we use an angled tip guidewire supported by a balloon catheter to penetrate the intima from the subintimal space thus gaining re-entry into a nonor less-diseased vessel segment as has been reported earlier for femoro-popliteal revascularization procedures [12]. Support catheters are not used. The intraluminal position is confirmed if the guidewire from below and another one inserted from above or a selective catheter touch each other as controlled by fluoroscopy. After that, we try to steer the guidewire from below into a selective catheter which has been pushed down as far as possible, and guide the wire out through the femoral sheath. If that fails, we push the guidewire from below all the way up to the groin and snare it in the proximal SFA from the femoral access, again hereby guiding the wire out. We then convert to an antegrade procedure by inserting a balloon catheter of the appropriate size and length via the femoral sheath to treat the BTK occlusion. If there are difficulties in traversing the lesion with the balloon, the guidewire is supported by pulling on both ends (pull-and-pull). Once the balloon is inflated within the lesion, the pedal sheath is immediately removed. We keep the balloon inflated for 120 s. After deflation of the balloon, the pedal access site is checked for hemostasis. If hemostasis is not achieved, manual compression may be applied by an assistant as needed while the procedure continues. After final angiogram via a selective catheter has confirmed technically successful revascularization (Fig. 3), the intervention is finished by removing the femoral



Fig. 3 A, B Via the femoral sheath a long conventional balloon is placed across the occlusion which is then dilated, starting in the distal segment of the occlusion. **C, D** DSA post POBA confirms sufficient revascularization of the anterior tibial artery. An intact access site within the dorsal pedal artery (white circle) is demonstrated by the final run-off angiogram.

Abb. 3 A, B Über die femorale Schleuse wird ein konventioneller PTA-Ballon eingewechselt und distal beginnend in der Läsion inflatiert. **C, D** die DSA nach der PTA zeigt eine suffizient revaskularisierte A. tib. anterior. Die retrograde Punktionsstelle der A. dorsalis pedis (weißer Kreis) stellt sich im Abschlussangiogramm unauffällig dar.

sheath and by using a vascular closure device or manual compression at the discretion of the operator.

Endpoints and definitions

Technical success was defined as intravascular placement of the pedal access vascular sheath within the target vessel after retrograde puncture of the dorsal pedal or distal tibial arteries as confirmed by manual retrograde angiogram via the sheath.

Crossing success was defined as successful retrograde passing of the target lesion intra- or subintimally with the guidewire tip in an intraluminal position above the target lesion.

Procedural success was defined as residual stenosis of less than $30\,\%$ following POBA.

Clinical success was defined as freedom from major amputation (limb salvage). Major amputation was defined as limb loss below or above the knee level, while minor amputation was defined as an amputation at the trans-metatarsal level or distal thereto. Procedure time was defined as the time from the first image

acquired to the last. The diameter (mm) of the retrogradely accessed pedal vessels was determined by a) calibrated measurements on angiographic images from the initial run-off angiogram (n=12) or b) by B-mode ultrasound images in axial orientation (n=4).

Treatment complications were categorized on the basis of outcome according to the reporting standards of the Society of Interventional Radiology [13]. There were two categories of minor complications: those resulting in (A) no therapy and no consequence or (B) nominal therapy and no consequence including overnight admission for observation only. Major complications included four categories: those (C) requiring therapy, minor hospitalization (48 h); (D) requiring major therapy, unplanned increase in level of care, prolonged hospitalization (48 h); (E) resulting in permanent adverse sequelae; and (F) resulting in death.

Statistical analysis

Data collection was performed by Excel (Microsoft). Statistical analysis was performed using commercially available software (SPSS, Version 22.0.0.0, IBM, USA). Continuous variables were expressed as mean +/- standard deviation and range. Kaplan-Meier analysis was used to predict the overall and major amputation-free survival. Probability values lower than 0.05 were considered to be significant.

Results



The mean follow-up was 197.8 days. The mean dose area product (DAP) was $2048.6cGy^*cm^2$. The mean procedure time was 92.4 + /-23 min. The retrogradely accessed vessel showed calcification of the punctured segment in 5/16 patients (31.3%). The mean diameter of the accessed vessels was 1.75 + /-0.24 mm.

The technical success rate was 100%. Crossing the lesion with the guidewire tip in an intra-luminal position from below was feasible in 12/16 patients (crossing success 75.0%). In 4 patients it was not possible to achieve re-entry. Further treatment in those 4 patients was conservative (1 patient refused surgery on the grounds of religious beliefs, and 3 patients lacked a suitable conduit or distal target vessel for bypass surgery). In 1 of the 12 patients the retrograde passage was subintimal, while the others were intraluminal. In 2 of these 12 patients the residual stenosis was > 30% following POBA and therefore procedural success was not achieved (procedural success in 10/16 patients (68.8%) and in 10/12 patients in which the lesion had been successfully traversed (83.3%)), although antegrade flow was documented.

The guidewire from below was guided out via the femoral sheath using a transfemoral snare maneuver in 8/12 interventions (66.7%), while the wire was steered into a diagnostic or balloon catheter inserted from above in 4/12 patients (33.3%). Prior to retrograde access of a crural vessel, 3/16 patients (18.8%) had successful revascularization of a femoro-popliteal lesion in the target limb during the same session. Hemostasis at the femoral access was achieved by manual compression in 14/16 patients (87.5%) and by a vascular closure device in 2/16 patients (12.5%). Manual compression of the pedal access site was performed in 4/16 patients (25.0%) for less than 5 minutes, while in the remaining patients hemostasis was found immediately after the PTA balloon in the target lesion had been deflated.

7 of 16 patients (43.8%) had undergone an ipsilateral minor amputation prior to retrograde revascularization. 1 patient (6.3%) had already had a major amputation of the contralateral limb. At baseline 7/16 patients (43.8%) had no patent BTK vessel, 8/16 patients (50.0%) had 1 and 1/16 patients (6.3%) had 2 patent crural vessels. At completion of the procedure, 2/16 patients (12.5%) had no, 5/16 (31.3%) had 1, 8/16 had 2 (50.0%) and 1/16 (6.3%) had 3 patent BTK vessels.

Minor complications occurred in 2/16 patients (12.5%) including one hematoma and vasospasm at the pedal access site in the other patient, both of which were treated conservatively. At completion angiography, all retrogradely accessed vessels were intact and showed no stenosis or occlusion at the point of access. Major complications were not observed.

The rate of freedom from major amputation (limb salvage) at 6 and 12 months was 87.5% and 72.9%, respectively (two major amputations on day 167 and 240). These two major amputations were not thought to be related to the transpedal access maneuvers but to a deterioration of ulcerations. One of these two patients with a major amputation had a failed BTK revascularization from antegrade as well from retrograde via the transpedal access. Both patients were in Rutherford-Becker class 6 at the time of revascularization. The survival rate at 6 and 12 months was 100% with one death due to pneumonia 18 months after revascularization (o Fig. 4A, B). Minor amputation after revascularization was observed in 1/16 patients (6.3%).

Discussion



Critical limb ischemia is associated with significant morbidity and mortality, particularly in patients with end-stage renal disease, who are undergoing dialysis or are diabetic [1, 2]. One major goal of treatment is to prevent (major) amputation in these pa-

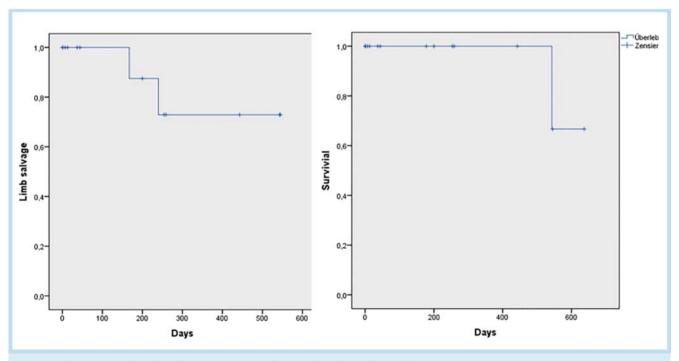


Fig. 4 Kaplan-Meier survival analysis of A major amputation-free survival and B overall survival.

Abb. 4 Kaplan-Meier-Analyse des A amputationsfreien und B Gesamtüberleben.

tients as it is thought to increase mortality rates [2]. Aggressive revascularization is indicated in CLI patients with tissue loss. The decision as to whether to revascularize by an endovascular approach or by open surgery may differ between centers with regard to expertise, lesion location and comorbidities. A recent study found that patients who are older and have more comorbidities were more likely to be treated by an endovascular approach than by lower extremity bypass. In this study patients who received endovascular therapy had lower perioperative mortality rates [14]. Therefore, in many centers a risk-adapted strategy is utilized with the endovascular approach representing the first-line therapy in older patients and those with relevant comorbidities, although popliteal-pedal bypass has shown better long-term patency rates [3]. Besides age and comorbidities with an increased operative risk, a lack of conduits and heavily calcified vessels may represent additional conditions in which one might favor an endovascular-first approach.

With modern equipment high technical success rates for BTK lesion revascularization have been reported. However, technical failure rates of up to 20% have been reported for the antegrade approach to BTK lesions [15]. Crossing the BTK lesion represents one major concern in the group with endovascular treatment failure [16]. Reasons for this might be a hard and fibrous proximal cap of the chronic total occlusion (CTO), unfavorably angled collaterals at the proximal end of the CTO as well as diffuse calcification [1, 8, 17]. In this context a retrograde approach was already described 25 years ago by surgeons who performed a cut-down of the posterior tibial artery [18]. Results after a percutaneous retrograde approach have recently been published with promising results regarding technical success and clinical outcomes [9, 11, 19, 20]. In the meantime a dedicated pedal access set has become commercially available. Walker et al. were the first to report a larger series of CLI patients who had been treated with this set and they reported a technical success rate of 95% for retrograde access to the dorsal pedal or distal tibial arteries [11]. Our rate of 100% technical success compares favorably with those and other published results.

In contrast to other working groups, we prefer to use ultrasound guidance with a linear probe for the retrograde access to the pedal or distal tibial arteries [5, 11]. We observed that the echogenic puncture needle included in the pedal access set can be sufficiently visualized by ultrasound. This might contribute to the high technical success rate observed in our study. Furthermore, radiation exposure of patients and staff is reduced. Even more importantly, in patients with often encountered end-stage renal disease the amount of contrast medium can be reduced by using ultrasound instead of angiography. However, in some patients it might be possible to visualize heavily calcified arteries and steer the puncture needle by use of fluoroscopy and without using contrast. In our experience a very low amount of subcutaneous tissue with a superficially located dorsal pedal artery or massive subcutaneous edema may be challenging scenarios and one option in such patients may be to relocate the puncture site more proximally if the distal extension of the target lesion allows for this.

There are three reasons why the retrograde passage of a BTK CTO might be successful where the antegrade has not been. First, the distal cap of the CTO is often softer and therefore easier to enter, especially if concave-shaped [8]. Second, from below the distance to the target lesion is shorter, and together with a smaller-sized access vessel, may give more support for the CTO wire and other equipment used. Third, collaterals do not misguide the wire as easily if coming from below owing to the angle and caliber of col-

laterals at the proximal end of the occlusion [1]. In fact, in selected cases careful analysis of the proximal (flat or convex) and distal cap (concave) of a CTO as well as existing collaterals may one day direct interventionalists to a retrograde-first approach. More data on the configuration of CTOs and how to tackle them best would be helpful. Regarding crossing the occlusion with a balloon, the second access at the pedal site allows for a pull-and-pull maneuver, thereby increasing the stability of the wire and enabling forward movement of the balloon which might have not been possible by an antegrade approach only.

Besides the technique of ultrasound-guided puncture, we altered the reported techniques for retrograde pedal interventions described by others in such a way that we remove the pedal access sheath as soon as we have converted to an antegrade maneuver, the target lesion has been crossed with an adequately sized balloon from above and the balloon has been inflated. In this way, antegrade blood flow to the pedal puncture site is reduced ("internal compression") and additional manual compression is hardly necessary. This may save time and increase the patients' comfort.

As an alternative to cross a BTK occlusion from above after conventional CTO guidewires have failed, dedicated crossing and reentry devices have become available. These devices may be used during the same session immediately after an antegrade wire passage of a lesion has failed and without any preparation for an alternative pedal access. Therefore, utilization of these devices might save time and be more comfortable for the patient compared to pedal puncture. As these devices add considerable costs to the procedure and are not reimbursed in the authors' country, we do not use these devices. Furthermore, reported procedural success rates for these devices are in the range of 70% [16], and therefore do not guarantee lesion crossing.

Regarding costs, the pedal access set and sterile dressing for the ultrasound probe not only need to be taken into account but the utilization of a snare device might also become necessary. To avoid utilization of a cost-adding snare device, we find it useful to first try to navigate the wire from below into a selective diagnostic or balloon catheter inserted from above. We try to place those as close to the lesion as possible to occlude as much as possible of the artery diameter, thereby enabling entry of the wire into the catheter and exit via the femoral sheath. However, as demonstrated in our study, this was technically successful in only one third of cases. Alternatively, the procedure may be finished from the pedal or tibial puncture site. Therefore, either a conventional 4F sheath has to be placed into a small-luminal vessel with the drawbacks of additional costs, risks of bleeding and the potential to induce spasm, or the procedure might be carried out in a retrograde manner using dedicated balloons which can be advanced through the pedal access set sheath. Furthermore, retrograde revascularization may be performed without a sheath by pushing the balloon through the arteriotomy with the risk of damage to the puncture site, especially during removal of the (partially) deflated balloon. We have limited experience with these alternatives and so far, for safety reasons, try to switch to an antegrade approach whenever possible.

The main reason for procedural failure in our study was inability to achieve re-entry after retrograde passage of the CTO. In this context one may hypothesize that the crossing and procedural success rates could be improved by use of additional techniques like the dual-balloon technique (balloons with antegrade and retrograde placement in the subintimal space with simultaneous inflation to interrupt the intimal wall) or by utilization of angulated (support) catheters [15]. However, during the study period

we did not use these techniques and did not have support catheters available. In contrast to our experience, Mustapha et al. reported the main reason for failure as access site calcification and consecutive inability to puncture the crural or pedal vessel [20]. Sabri et al. reported both an inability to access the pedal vessel and failure to achieve re-entry following retrograde subintimal passing of the lesion as the main reasons for procedural failure [19].

So far, the literature regarding retrograde transpedal revascularization procedures is limited and to some degree inhomogeneous. Walker et al. [11] reported the largest series of patients. They too found a high technical success rate of 95 %. However, in their cohort also patients with femoral and/or popliteal occlusions were treated by a retrograde approach, making it difficult to compare it with our group with BTK occlusions only. Furthermore, their definition of procedural success was "antegrade flow" in the target lesion, while we chose a more rigorous endpoint for the definition of procedural success with residual stenosis <30%. This might explain the higher success rate in their study (99% vs. 69%). Unfortunately, they did not report on limb salvage rates. Other groups, however, reported limb salvage rates at 12 months between 64% and 82.3% after retrograde pedal access interventions [1, 5, 19]. Although femoral and popliteal occlusions were included in these studies, the reported limb salvage of 72.9% in our study compares favorably with these results.

Compared to the results reported by Bazan et al. [1], we noted with interest that although the lesions in our study appear to be more complex (only CTOs, longer lesions), the results for procedural success and limb salvage are similar. From this we conclude that the complexity of a lesion may play a minor role regarding outcome as long as it can be passed with a guidewire from the retrograde approach.

Possible drawbacks of the retrograde access maneuver may include secondary changes like hematoma, scar formation and vascular injuries at the pedal access site which may complicate a pedal or tibial bypass. However, regarding complications our results are comparable to those reported by others and underline the fact that the retrograde pedal or tibial access is safe. Walker et al. [11] reported one pedal occlusion at the access site (0.4%), of note after use of a 6F sheath. Therefore, one may anticipate that a retrograde puncture does not preclude a pedal bypass. Other reported complications relate mainly to ones observed at the femoral access (hematoma) and myocardial infarction. Major complications have been reported in up to 8 % of cases, and minor complications in up to 2% [1, 19]. We observed no major complications and found a minor complication rate of 12.5 %. Taking into account that major complications are mainly associated with the femoral access, maybe more procedures should be performed solely from the retrograde access. However, in such cases careful, high-quality, noninvasive imaging is mandatory to evaluate inflow lesions and plan the retrograde approach. On the other hand, in our experience two or even all crural vessels often need to be treated during the intervention and an exclusively retrograde approach may be time-consuming or even impossible in such a scenario but may be a good option if only one vessel has to be treated.

There are some limitations to this study. First, the number of patients is small, which prevents us from generalizing our data. Second, the presented series is retrospective and lacks randomization. Therefore, patient selection bias may have influenced the results. Furthermore, no rigorous imaging follow-up was available in the analyzed patients. Therefore, no data on patency or

target lesion revascularization of the lesions with retrograde revascularization are available.

In conclusion, the retrograde transpedal or transtibial access utilizing a dedicated pedal access set is safe and offers high technical and acceptable clinical success rates. BTK occlusions that could not be addressed by an antegrade transfemoral approach should be tackled by a retrograde approach whenever possible. Further data are needed to answer the question as to whether this could be the first-line approach in selected patients as compared to the antegrade transfemoral route.

All procedures performed in this study were in accordance with the ethical standards of the institutional research committee and with the 1964 Helsinki declaration and its later amendments. The ethics approval number for this study is 15 – 285A.

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