

Review Article

Countermeasures against Infection in Critical Limb Ischemia Treatments

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The preoperative and postoperative infection control measures for critical limb ischemia treatments were described. The treatment strategies for severe ischemic limbs were showed according to the presence and extent of infection. If the treatment strategy for a severe ischemic limb with infection is mistaken, infection will spread and make worse the situation of the ischemic limb, and eventually it can result not only in limb loss but also life threatening. A surgical strategy is very important in the bypass material, the selection of anastomotic site, the use of postoperative antibacterial drugs, and the wound treatment. Infection troubles are the most familiar and indispensable problem for surgeons, the countermeasures against infection especially in critical limb ischemia is the key point along with revascularization. (This is a translation of *Jpn J Vasc Surg* 2018; 27: 129–132.)


Keywords: critical limb ischemia, infection, bypass, surgical site infection

Introduction

A reliable revascularization is crucial for the treatment of critical limb ischemia (CLI) as the outcome can be significantly affected by concomitant infections of ulcers and necrotic sites. Concomitant diabetes is particularly prone to complicating infections prior to surgery.¹⁾ In this paper, we discuss CLI treatment strategies, including pre- and postoperative infection control.

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Evaluation of Limb Ischemia

Effective CLI treatment relies on a proper evaluation of the extent of ischemia, the presence or absence of concomitant infections and their degree, and prompt treatment initiation. The extent of ischemia is determined from subjective symptoms, ankle–brachial pressure index, skin perfusion pressure, and size of any skin ulcers or tissue defects. Prompt evaluation of the presence or absence and spread of concomitant infections is also necessary. Even if an infection appears to be confined to a certain area, it should be carefully monitored for unexpected progression and growth of abscesses along the tendons and ligaments and towards the proximal direction of the foot.²⁾ This is related to the anatomical structure of the foot, which is divided into three fascial cavities, inside which blood vessels travel in a mesh pattern (Fig. 1). Once infection spreads to one of these fascial cavities, drainage becomes difficult, facilitating the longitudinal progression (proximal progression) of the lesions (Fig. 2). In some cases, an adequate drainage can only be achieved through metatarsal amputation.

Moreover, this condition often becomes severe in patients with poorly controlled diabetes. It is therefore crucial to manage risk by monitoring the general condition alongside local evaluation. Furthermore, deep ulcers present at the region where the bone is exposed can further complicate osteomyelitis and arthritis, thus requiring confirmation of bone changes through magnetic resonance imaging and computed tomography in addition to angiography.³⁾

Treatment Strategy

Figure 3 shows a CLI treatment strategy that accounts for the presence or absence of concomitant infection as well as its degree.

In the absence of an infection, CLI treatment aims to preserve the limb and improve the patient's quality of life by performing the minimum required systemic evaluation followed by revascularization. In the presence of an infection, which is confined to one area, the treatment will comprise either revascularization followed by debride-

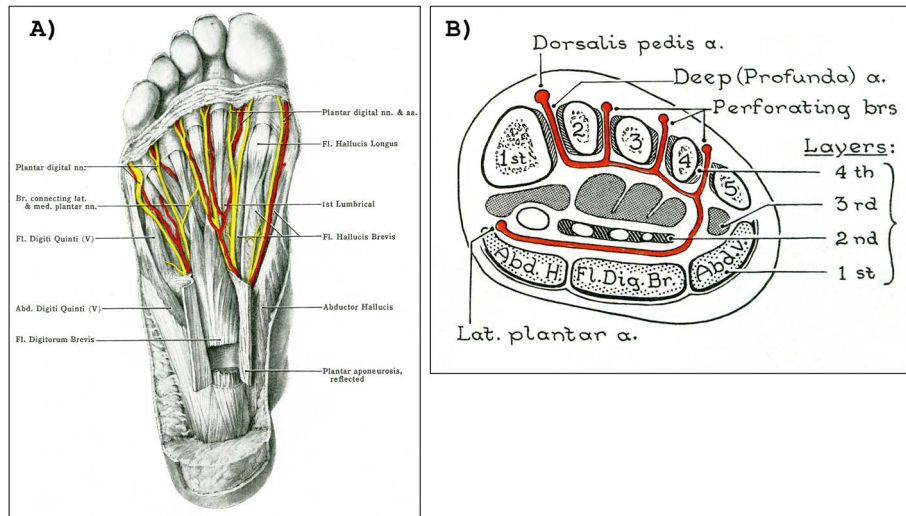


Fig. 1 A) Anatomy of plantar muscles, digital nerves and arteries. B) Cross-section near metatarsal bases. (From Anderson: *Grant's Atlas of Anatomy*. 8th ed.).



Fig. 2 Infection expanded into plantar fascia space.

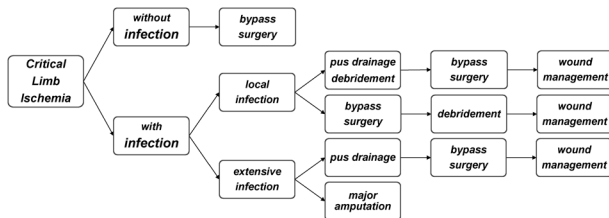


Fig. 3 Treatment strategy of clinical limb ischemia.

ment or drainage and debridement followed by revascularization. The latter can favor the spread of necrosis and infection, hence exacerbating limb ischemia if the prospect for revascularization time is missed, resulting in the need for excessive debridement. A treatment plan therefore must be devised promptly after diagnosis. It is important to keep in mind that preventing the spread of infection can minimize tissue defects.

Even in patients with spreading infections, the limb can be preserved to some extent through proper preoperative drainage and wound treatment followed by reliable revascularization. However, investigation into the selection of anastomotic sites is needed. Extensive infection that spreads further into the proximal direction of the

ankle joint and CLI that causes cellulitis in the lower leg have an extremely poor prognosis and can require major amputation to save the patient's life without performing revascularization.

The recommended treatment for infection is broad-spectrum antibiotics formulated with carbapenems or beta-lactamase inhibitors.⁴⁾ Anaerobic bacteria are particularly prone to being isolated from the deep tissue of ulcers in patients with concomitant diabetes.⁵⁾ Cases of mixed infection of methicillin-resistant *Staphylococcus aureus* and *Pseudomonas aeruginosa* can also be observed. While no consensus has been reached regarding the systemic administration of antibiotics to patients with local wound infection and no systemic signs of infection,⁶⁾ it is recommended to select an antibiotic that acts on the bacterium in question throughout the pre- and postoperative periods in patients where the wound bacterium has been identified preoperatively.⁷⁾

Revascularization and Surgical Site Infection Control

Guidelines for the proper use of antibiotics as postoperative infection prophylaxis, including surgical site infection (SSI) control, have been created by the Japanese Society of Chemotherapy and the Japan Society for Surgical Infection. For the specific case of vascular surgery, guidelines for the use of antibiotics to prevent postoperative infection are presented in Table 1.⁷⁾ In vascular surgery, prophylactic antibiotics target resident skin bacteria such as *S. aureus* and Streptococci, thus cefazolin and sulbactam/ampicillin are commonly selected for SSI control. These antibiotics are administered within 1 h prior to skin incision and re-administered if surgery is prolonged. Re-

Table 1 Perioperative prophylactic anti-biotics usage in vascular surgery (the abstract from ref. 7)

Surgical procedure	Recommendation grade/evidence level for indication	Recommended antibiotic	Alternative drugs for patients with allergies	Administration period		Remarks
				Single dose or postoperative time	Recommendation grade/evidence level	
Surgery with artificial prosthesis: abdominal aortic surgery	A-1	CEZ	VCM, CLDM	24–48 h	C1-III	Elective surgery should be performed after infection control through antibacterial treatment of infected aneurysm (C1-III)
Surgery with artificial prosthesis: Abdominal aortic surgery (high risk of SSI)	A-1	CEZ	VCM, CLDM	72 h	C1-III	*High risk of SSI: emergency surgery for rupture, etc.
Surgery with artificial prosthesis: lower limb bypass surgery	A-1	CEZ	VCM, CLDM	Single dose–24 h	A-1	
Endovascular treatment for peripheral blood vessels	C1-III	CEZ	CLDM, VCM	Single dose	C1-III	
Lower limb revascularization (no use of artificial prosthesis)	C1-III	CEZ	CLDM, VCM	Single dose (re-administration after a long period of time)	C1-III	Selection of an antibiotic susceptible to isolates in the event of lower limb ulceration, etc. (C1-III)

CEZ: cefazolin, VCM: vancomycin, CLDM: clindamycin, SSI: surgical site infection.

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Fig. 4 NPWT (Negative Pressure Wound Therapy) after digital amputation.

administration is generally recommended at intervals of two-fold of the half-life.⁷⁾

Whenever possible, revascularization for CLI should be

performed through the use of a vein graft, making every effort to avoid the use of artificial blood vessels. In patients who require reconstruction of multiple segments, a distal bypass is commonly employed avoiding surgical invasion to the inguinal region as much as possible, where the lymph nodes and vessels concentrate and infection is common. This bypass should be combined with endovascular treatment in the regions of the iliac artery and superficial femoral artery. If an artificial blood vessel is required for an extra-anatomical bypass, expanded polytetrafluoroethylene, which is hydrophobic and presents low histocompatibility and porosity, is advantageous.⁸⁾ However, the incidence of infection is high in extra-anatomical bypass, redo surgery, concomitant diabetes, and CLI.⁹⁾ Furthermore, old age, obesity, steroid and immunosuppressant use, emergency surgery, prolonged surgery, and other factors pose a high risk for the onset of SSI.⁷⁾ In particular, graft sepsis following wound infection can have fatal consequences.¹⁰⁾ Therefore, full consideration must be given when selecting the treatment strategy, surgical procedure,

and graft. The risk of postoperative wound infection is high in cases of foot ulcers and necrosis, indicating that caution must be paid to the site of peripheral anastomosis during distal bypass surgery. Even if revascularization is successful and blood flow is ensured, infection of an anastomotic wound can lead to rupture of the anastomosis and ultimate loss of the limb.

Wound Treatment and Management

Contaminants and deposits on the wound surface inhibit healing. Local wound cleaning is therefore crucial in removing such contaminants and deposits when treating tissue defects. From the infection viewpoint, the wound cleaning solution used (physiological saline, acidic water, tap water, etc.) does not affect the outcome.¹¹⁾ The simple use of a topical antibiotic can generate resistant bacteria and thus should be carefully considered.

Once the blood flow has been ensured and the infection has been controlled to a certain degree, treatment to promote granulation is required. In 1997, Morykwas et al.¹²⁾ reported the usefulness of negative-pressure wound therapy (NPWT), after which this practice became widespread for the treatment of refractory wounds. In Japan, from 2010, NPWT is only applied under health insurance coverage and has led to good outcomes when applied to various wounds (Fig. 4).

Conclusion

This paper discussed pre- and postoperative infection control in terms of CLI treatment. For surgeons, infection is the most common and familiar complication and in the specific case of CLI, it is critical for successful treatment alongside revascularization.

Disclosure Statement

There are no conflicts of interest.

Additional Note

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