Upper Limb Ischemia: Clinical Experiences of Acute and Chronic Upper Limb Ischemia in a Single Center

Miju Bae, M.D., Sung Woon Chung, M.D., Chung Won Lee, M.D., Jinseok Choi, M.D., Seunghwan Song, M.D., Sang-pil Kim, M.D.

**Background:** Upper limb ischemia is less common than lower limb ischemia, and relatively few cases have been reported. This paper reviews the epidemiology, etiology, and clinical characteristics of upper limb ischemia and analyzes the factors affecting functional sequelae after treatment. **Methods:** The records of 35 patients with acute and chronic upper limb ischemia who underwent treatment from January 2007 to December 2012 were retrospectively reviewed. **Results:** The median age was 55.03 years, and the number of male patients was 24 (68.6%). The most common etiology was embolism of cardiac origin, followed by thrombosis with secondary trauma, and the brachial artery was the most common location for a lesion causing obstruction. Computed tomography angiography was the first-line diagnostic tool in our center. Twenty-eight operations were performed, and conservative therapy was implemented in seven cases. Five deaths (14.3%) occurred during follow-up. Twenty patients (57.1%) complained of functional sequelae after treatment. Functional sequelae were found to be more likely in patients with a longer duration of symptoms (odds ratio, 1.251; \( p = 0.046 \)) and higher lactate dehydrogenase (LDH) levels (odds ratio, 1.001; \( p = 0.031 \)). **Conclusion:** An increased duration of symptoms and higher initial serum LDH levels were associated with the more frequent occurrence of functional sequelae. The prognosis of upper limb ischemia is associated with prompt and proper treatment and can also be predicted by initial serum LDH levels.

Key words: 1. Upper extremity  
2. Ischemia  
3. Complication  
4. L-lactate dehydrogenase

**INTRODUCTION**

Upper limb ischemia is less common than lower limb ischemia, and relatively few cases have been reported. However, delays in diagnosis and treatment are likely to result in severe functional impairment and disability, even in the absence of overt tissue loss [1,2]. In a clinical context, the shoulder and elbow are much more tolerant of ischemia due to their well-developed collateral circulation, and it is therefore more common to observe ischemic symptoms below the elbow [3]. This study presents a review of the epidemiology, etiology, and clinical characteristics of upper limb ischemia, as well as an analysis of the factors affecting functional sequelae after treatment.
Clinical Experiences of Acute and Chronic Upper Limb Ischemia in a Single Center

Table 1. Demographic characteristics of the patients

| Characteristic          | Number (%) |
|------------------------|------------|
| Gender                 |            |
| Male                   | 24 (68.6)  |
| Female                 | 11 (31.4)  |
| Heart disease          | 18 (51.4)  |
| Atrial fibrillation    | 9 (25.7)   |
| Valvular disease       | 3 (8.6)    |
| Ischemic disease       | 2 (5.7)    |
| Congestive heart failure| 2 (5.7)   |
| Bradycardia            | 1 (2.9)    |
| Dilated cardiomyopathy | 1 (2.9)    |
| Hypertension           | 15 (42.9)  |
| Stroke                 | 6 (17.1)   |
| Multiple trauma        | 5 (14.3)   |
| Diabetes mellitus      | 3 (8.6)    |
| Rheumatic arthritis    | 2 (5.7)    |
| Systemic scleroderma   | 2 (5.7)    |

Table 2. Diagnostic tools

| Diagnostic tools                        | Number (%) |
|-----------------------------------------|------------|
| CT angiography                          | 22 (62.9)  |
| Conventional angiography                | 10 (28.6)  |
| Only history taking and physical examination | 2 (5.7)  |
| Duplex ultrasound                       | 1 (2.9)    |

Table 3. Etiology

| Etiology                      | Number (%) |
|-------------------------------|------------|
| Acute                         | 19         |
| Embolus from heart origin     | 11 (31.4)  |
| Trauma                        | 7 (20.0)   |
| Embolus from aortic arch      | 1 (2.9)    |
| Chronic                       | 16         |
| Raynaud’s disease             | 4 (11.4)   |
| Burger’s disease              | 3 (8.6)    |
| Atherosclerosis               | 2 (5.7)    |
| Thoracic outlet syndrome      | 2 (5.7)    |
| Unknown                       | 5 (14.3)   |
| Total                         | 35         |

Table 4. Locations of lesions

| Locations                        | Number (%) |
|----------------------------------|------------|
| Innominate artery                | 1 (2.9)    |
| Subclavian artery                | 4 (11.4)   |
| Axillary artery                  | 3 (8.6)    |
| Brachial artery                  | 17 (48.6)  |
| Radial artery                    | 2 (5.7)    |
| Ulnar artery                     | 4 (11.4)   |
| Palmar arterial arch             | 4 (11.4)   |

RESULTS

A total of 35 patients who underwent treatment for acute and chronic upper limb ischemia in a single center between January 2007 and December 2013 were reviewed. Cases involving arterial insufficiency after the creation of arteriovenous fistulae for hemodialysis were excluded. The baseline characteristics, comorbidities, etiologies, diagnostic tools, locations of the lesion, treatments, complications, and sequelae after treatment were reviewed. Additionally, the factors that were expected to affect to functional sequelae after treatment were analyzed using binary logistic regression. The results were expressed as means with 95% confidence intervals where appropriate, and p-values < 0.05 were considered to imply statistical significance. Statistical analysis was performed using IBM SPSS ver. 20.0 (IBM Co., Armonk, NY, USA).

The median age of the patients was 55.03 years, and the median follow-up duration was 706.23 days (23.54 months). Fifteen patients (42.9%) were current smokers and four (11.4%) were ex-smokers. Table 1 summarizes the demographic characteristics of the patients in this study.

The initial diagnosis of upper limb ischemia was mostly made on the basis of computed tomography (CT) angiography (62.9%). Conventional angiography and duplex ultrasound were also used as diagnostic tools. Two cases were diagnosed solely on the basis of a medical history and physical examination (Table 2). The most common etiology was embolism of cardiac origin (31.4%), followed by thrombosis with secondary trauma (20.0%) (Table 3), and the brachial artery (48.6%) was the most common location of lesions causing obstruction (Table 4).

A total of 28 operations were performed, while seven patients underwent conservative therapy (Table 5). The oper-
Table 5. Treatments

| Treatment                                      | Number (%) |
|------------------------------------------------|------------|
| Embolectomy or thrombectomy                    | 16 (45.7%) |
| Bypass with great saphenous vein               | 5 (14.3%)  |
| Percutaneous catheter direct thrombolysis      | 4 (11.4%)  |
| Primary repair                                  | 2 (5.7%)   |
| Sympathectomy                                   | 1 (2.9%)   |
| Conservative therapy                            | 7 (20.0%)  |

Table 6. Complications after treatment

| Complications          | Number (%) |
|------------------------|------------|
| Re-obstruction         | 2 (5.7%)   |
| Wound infection        | 2 (5.7%)   |
| Acute renal failure    | 2 (5.7%)   |
| Minor amputation       | 2 (5.7%)   |
| Bleeding               | 1 (2.9%)   |

In the case of embolism with atrial fibrillation, emergency embolectomy was performed first, followed by echocardiography to re-evaluate the status of the heart. Isolated atrial fibrillation was treated with anticoagulation drugs. If a patient had a history of other heart diseases, the treatment strategy was determined in consultation with the cardiology department. Imaging of the brain, lower extremities, or mesenteries was not performed if a patient did not complain of symptoms in other areas.

Treatment-related complications are described in Table 6. Additionally, five deaths (14.3%) occurred during the course of follow-up. One of those deaths (2.9%) occurred within 30 days of the operation, and was due to postoperative rhabdomyolysis with acute renal failure. The other deaths were due to acute small intestinal infarction (1), acute myocardial infarction (1), and unknown causes (2).

Two cases of reobstruction were noted. One case occurred seven months after surgery, and the other occurred four years after surgery. In each of these patients, reobstruction occurred in the same location as the previous lesion, and percutaneous catheter-directed thrombolysis was performed.

Functional sequelae were observed in 20 patients (57.1%), with different patterns of symptoms in each patient. The sequelae after treatment included decreased motor function (25.7%), decreased sensory function (14.3%), persistent pain (5.7%), and tingling sensations in the fingers or on the palm (40%).

Factors thought to possibly affect the emergence of sequelae were analyzed (Table 7). Functional sequelae were found to be significantly more common in patients with a longer duration of symptoms (p=0.046) and higher initial serum lactate dehydrogenase (LDH) levels (p=0.031). The mean initial serum LDH level of the patients who experienced functional sequelae was 454.71 IU/L, which was significantly higher than the normal upper reference limit of 225 IU/L.

Table 7. Result of binary logistic regression for functional sequelae

| Variable                                      | p-value | Odds ratio |
|-----------------------------------------------|---------|------------|
| Gender                                        | 0.348   | 0.278      |
| Age                                           | 0.531   | 1.026      |
| Symptom duration                              | 0.046   | 1.521      |
| Hypertension                                  | 0.959   | 0.939      |
| Diabetes                                      | 0.698   | 1.964      |
| Smoking                                       | 0.725   | 1.372      |
| Atrial fibrillation                           | 0.582   | 0.380      |
| Coronary arterial occlusive disease           | 0.999   | 1.885      |
| Stroke                                        | 0.343   | 4.230      |
| Hemoglobin                                    | 0.742   | 1.202      |
| Platelet                                      | 0.992   | 0.943      |
| White blood cell                              | 0.841   | 0.875      |
| Aspartate aminotransferase                    | 0.771   | 0.982      |
| Alanine aminotransferase                      | 0.802   | 1.002      |
| Lactate dehydrogenase                         | 0.031   | 1.001      |
| Albumin                                       | 0.089   | 0.724      |
| Phosphate                                     | 0.425   | 1.259      |
| C-reactive protein                            | 0.074   | 1.424      |
| Creatine kinase                               | 0.885   | 1.576      |
| Myoglobin                                      | 0.647   | 1.059      |

DISCUSSION

Upper limb arterial ischemia is responsible for <5% of all cases of limb ischemia [4]. The epidemiology of upper limb ischemia...
Clinical Experiences of Acute and Chronic Upper Limb Ischemia in a Single Center

arterial disease is difficult to assess, due to its low incidence and vast number of etiologies [5]. The anatomical location and etiology are the two major criteria for categorizing upper limb arterial disease. The anatomical location is subdivided into the small or large arteries, and the etiology is subdivided into either occlusive disease or vasospasm [5].

The diseases that lead to the most severe cases of upper limb arterial ischemia are autoimmune or connective tissue diseases such as scleroderma, rheumatoid arthritis, systemic lupus, and others. Although Burger’s disease (thromboangiitis obliterans) most commonly affects the lower limbs, approximately 50% of patients also have upper limb involvement with subsequent digital ischemia [6].

The most common large-vessel arteriopathy in the upper limb is atherosclerosis [7], and the most common occlusive site is the origin of the left subclavian artery. Embolism is the most common cause of acute upper limb ischemia [8]. Most emboli originate from the heart, caused by atrial fibrillation, recent myocardial infarction, and valvular heart disease [9]. Embolic occlusion from atrial fibrillation or other sources is classically seen in the brachial artery before the bifurcation of the radial and ulnar arteries [5].

In this study, the most common cause of upper limb ischemia was found to be embolism, and the distal brachial artery was observed to be the most frequent site of obstructions (Tables 3, 4). In this study, 19 patients (54.3%) had acute limb ischemia. The mean age of patients with acute upper limb ischemia was 60.42 years, and female patients comprised 42.10% of the study population.

Spinelli et al. [10] reported that duplex ultrasound was the most widely used diagnostic tool. In contrast, Licht et al. [3] reported that 88% of operations were performed based only on the patient’s medical history and a physical examination. In our center, when upper limb ischemia was suspected from a patient’s medical history and a physical examination, CT angiography was used as the initial diagnostic tool in 62.9% of cases. CT was useful in locating the lesion, discovering multiple lesions, and assessing the correlation of the lesion with the surrounding structure. Between 9% and 30% of patients with upper limb arterial occlusive disease seen by vascular surgeons are managed nonoperatively due to significant comorbidities or minimal symptomatology [11].

However, the treatment of choice for acute upper limb ischemia is surgical correction via embolectomy using a Fogarty balloon catheter. Incision of the antecubital fossa still remains the best treatment for removing brachial emboli [9].

Revascularization with upper limb bypass is used less commonly than other treatments, comprising approximately 4% of all vascular operations [12]. Although this procedure is infrequently performed, the results of upper extremity bypass are excellent, and are even superior to those reported for the corresponding treatment of lower extremity ischemia [5]. Brunkwall et al. [13] reported a postoperative two-year recanalization rate of 60%–90%. Moreover, Spinelli et al. [10] reported a primary patency rate of 82.6% and a secondary patency rate of 91.3% over a 34-month follow-up period [9]. In our study, 23 patients (65.7%) underwent surgical correction, of whom five (14.3%) underwent bypass surgery using the reversed great saphenous vein.

Recently, the use of endovascular treatment to treat various vascular diseases has grown exponentially. Kim et al. [14] performed percutaneous aspiration thromboembolectomy and thrombolysis in 11 patients suffering from acute upper limb ischemia, of whom nine experienced successful recanalization. In this study, four patients who suffered from acute brachial artery embolism underwent percutaneous aspiration thromboembolectomy and thrombolysis, and one of those patients died due to rhabdomyolysis with acute renal failure. In endovascular treatment, the embolus or distal embolism is dissolved after percutaneous aspiration thromboembolectomy employing various techniques. An advantage of this technique is the absence of an operative wound. Moreover, the recovery time is fairly short. However, a disadvantage is that thrombolysis requires more time for revascularization and involves a longer ischemic time, and hemorrhagic complications are more frequent. Thus, endovascular treatment is unsuitable for the reperfusion of acute cases of critical upper limb ischemia, and should be limited to stage I and IIa acute limb ischemic lesions or the cautious treatment of chronic ischemia.

In this study, some cases of upper limb ischemia were associated with Raynaud’s phenomenon, which involves episodic vasospasm when the patient is exposed to cold or psychosocial stress. It usually resolves naturally, but can persist and become a problem. If the vasospasm continues, con-
Conservative management is implemented using medications such as calcium channel blockers and phosphodiesterase inhibitors. Surgical intervention is performed if the patient is unresponsive to conservative management. In this study, four patients suffered from Raynaud’s phenomenon; two patients had systemic scleroderma as the underlying disease, and the other two had rheumatic arthritis as the underlying disease. Two patients underwent the amputation of a digit due to the progression of finger necrosis. All four patients were prescribed calcium channel blockers and antiplatelet agents. Those who were unresponsive to medication were selectively treated with video-assisted thoracoscopic surgery and sympathectomy. Cervical sympathectomy is no longer considered an appropriate treatment for Raynaud’s phenomenon because the very short duration of its effectiveness. However, the long-term results of periar terial sympathectomy are more reliable than those of cervical sympathectomy [15].

Most patients were admitted to the emergency department with the acute onset of symptoms resulting from embolism and thrombosis. In such cases, the duration of symptoms is usually short. However, patients with Raynaud’s syndrome, Burger’s disease, or thoracic outlet syndrome-related upper limb ischemia visited through the outpatient clinic, and had a longer duration of symptoms. The duration of symptoms in cases of upper limb ischemia may vary from two hours to a year, depending on the etiology and severity of the illness. Many debates have addressed whether the time gap between the onset of symptoms and treatment predicts long-term arm function. Abbott et al. [16] reported that patients treated within 12 hours exhibited a lower mortality rate and a higher rate of limb salvage. Elliott et al. [17] also found that the results of treatment showed a linear relationship to the time gap between symptom onset and treatment. Bang and Nalachandran [18] also found that the prognosis differed significantly when >12 hours elapsed between the onset of symptoms and treatment. However, it has been suggested that the time gap does not influence the prognosis [3]. In this study, the time gap between the onset of symptoms and consequent treatment was related to postoperative sequelae. A longer duration of symptoms was associated with a higher likelihood of sequelae (odds ratio, 1.251; p=0.046).

In this study, serum LDH levels were also found to have a significant influence on the occurrence of functional sequelae (odds ratio, 1.001; p=0.031). Higher levels of serum LDH were associated with a higher chance of functional sequelae. LDH has been used as a marker for liver disease, heart attack, anemia, muscle trauma, bone fracture, cancer, and infections such as meningitis, encephalitis, and HIV; when tissue is damaged by injury and disease, LDH is released into the bloodstream. An initial increase in LDH reflects tissue damage resulting from upper limb ischemia, and initial tissue damage is related to postoperative functional sequelae. Levels of creatine kinase or the myoglobin enzymes with muscle specificity could theoretically lead to an accurate assessment of the prognosis, but were found to show no statistically significant associations in this study. Nonetheless, the use of serum LDH levels as a basic liver function test was able to predict the development of functional sequelae to a statistically significant extent.

In conclusion, a longer duration of symptoms and higher initial serum LDH levels were associated with a greater likelihood of functional sequelae. The prognosis of upper limb ischemia is associated with prompt and proper treatment and can also be predicted by initial serum LDH levels.

**CONFLICT OF INTEREST**

No potential conflict of interest relevant to this article was reported.

**ACKNOWLEDGMENTS**

This study was supported by a grant from the Pusan National University Hospital grant (2013).

**REFERENCES**

1. Galbraith K, Collin J, Morris PJ, Wood RF. Recent experience with arterial embolism of the limbs in a vascular unit. Ann R Coll Surg Engl 1985;67:30-3.
2. Pentti J, Salenius JP, Kuukasjarvi P, Tarkka M. Outcome of surgical treatment in acute upper limb ischaemia. Ann Chir Gynaecol 1995;84:25-8.
3. Licht PB, Balezantis T, Wolff B, Baudier JF, Roder OC. Long-term outcome following thrombembolectomy in the up-
Clinical Experiences of Acute and Chronic Upper Limb Ischemia in a Single Center

per extremity. Eur J Vasc Endovasc Surg 2004;28:508-12.
4. McCarthy WJ, Flinn WR, Yao JS, Williams LR, Bergan JJ. Result of bypass grafting for upper limb ischemia. J Vasc Surg 1986;3:741-6.
5. Alef MJ, Hamdan A. Upper extremity arterial disease: general considerations. In: Rutherford RB, editor. Vascular surgery. 8th ed. Philadelphia: Elsevier Saunders; 2014. p. 1868-74.
6. Olin JW. Thromboangiitis obliterans (Buerger’s disease). N Engl J Med 2000;343:864-9.
7. Campbell WB, Ridler BM, Szymanska TH. Current management of acute leg ischaemia: results of an audit by the Vascular Surgical Society of Great Britain and Ireland. Br J Surg 1998;85:1498-503.
8. Eyers P, Earnshaw JJ. Acute non-traumatic arm ischaemia. Br J Surg 1998;85:1340-6.
9. Deguara J, Ali T, Modarai B, Burnand KG. Upper limb ischaemia: 20 years experience from a single center. Vascular 2005;13:84-91.
10. Spinelli F, Benedetto F, Passari G, et al. Bypass surgery for the treatment of upper limb chronic ischaemia. Eur J Vasc Endovasc Surg 2010;39:165-70.
11. Roddy SP, Paty P. Upper extremity arterial disease: revascularization. In: Rutherford RB, editor. Vascular surgery. 8th ed. Philadelphia: Elsevier Saunders; 2014. p. 1875-84.
12. Bergqvist D, Ericsson BF, Konrad P, Bergentz SE. Arterial surgery of the upper extremity. World J Surg 1983;7:786-91.
13. Brunkwall J, Bergqvist D, Bergentz SE. Long-term results of arterial reconstruction of the upper extremity. Eur J Vasc Surg 1994;8:47-51.
14. Kim SK, Kwak HS, Chung GH, Han YM. Acute upper limb ischemia due to cardiac origin thromboembolism: the usefulness of percutaneous aspiration thromboembolectomy via a transbrachial approach. Korean J Radiol 2011;12:595-601.
15. Herrick AL, Muir L. Raynaud’s phenomenon. In: Rutherford RB, editor. Vascular surgery. 8th ed. Philadelphia: Elsevier Saunders; 2014. p. 1901-14.
16. Abbott WM, Maloney RD, McCabe CC, Lee CE, Wirthlin LS. Arterial embolism: a 44 year perspective. Am J Surg 1982;143:460-4.
17. Elliott JP Jr, Hageman JH, Szilagyi E, Ramakrishnan V, Bravo JJ, Smith RF. Arterial embolization: problems of source, multiplicity, recurrence, and delayed treatment. Surgery 1980;88:833-45.
18. Bang SL, Nalachandran S. Upper limb ischaemia: a single centre experience. Ann Acad Med Singapore 2009;38:891-3.