

Extracorporeal Shock Wave Therapy for Painful Chronic Neurogenic Heterotopic Ossification After Traumatic Brain Injury: A Case Report

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Neurogenic heterotopic ossification (NHO) is a process of benign bone formation and growth in soft tissues surrounding major synovial joints and is associated with central nervous system (CNS) injuries. It is a common complication in major CNS injuries, such as traumatic brain injury, spinal cord injury, and stroke. Here, we report the case of a 72-year-old male, who experienced a traumatic brain injury and painful chronic NHO around the left hip joint. Three applications of extracorporeal shock wave therapy (ESWT) were administered to the area of NHO, which resulted in pain relief and an improvement in the loss of motion in the left hip joint. Improvements were also noted in walking performance and activities of daily living, although the size of NHO remained unchanged. Therapeutic effects of ESWT lasted for 12 weeks.

Keywords Heterotopic ossification, Extracorporeal shock wave therapy, Brain injury

INTRODUCTION

Neurogenic heterotopic ossification (NHO) is a common complication in major central nervous system injuries, such as traumatic brain injury (TBI), spinal cord injury, and stroke [1]. It is a localized and progressive formation of pathological ectopic bone in soft tissues at the major synovial joints. Significant morbidity can occur at

the site of NHO, including pain and loss of motion (LOM), and can interfere with activities of daily living and reduce the quality of life. Therefore, treatment of NHO is important for improved rehabilitation.

Extracorporeal shockwave therapy (ESWT) has been shown to be an effective method of treating both acute and chronic soft-tissue painful inflammations [2]. The mechanism of ESWT is not fully understood, but it may work via anti-inflammatory mechanism [3] and tissue regeneration with neovascularization [4] and reduction of neurogenic inflammation [5].

We experienced a case with chronic painful NHO after TBI. As a mature NHO, intense inflammation was gone. Nevertheless, the severe pain had been preserved in the affected area and intractable. We thought soft tissues around NHO were compressed by calcifying mass and these mass-effects led to local ischemic injury and ne-

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crisis. Progression of NHO is enough to compress nearby neurovascular structures and thereby cause hypoxic tissue damage [6,7].

We hypothesized that the recovery of soft tissue injury by mass-effects around NHO can improve the intractable pain and expected ESWT would be a useful method to enhance tissue regeneration via several mechanisms mentioned above.

To the best of our knowledge, there are no prior reports regarding the use of ESWT as a treatment for chronic painful NHO in Asian patients. We herein report this case with a review of the literature.

CASE REPORT

A 72-year-old male with no notable past medical history except diabetes and hypertension was in a pedestrian traffic accident on October 13, 2012. Craniectomy and cranioplasty were performed to treat the resulting TBI. In June 2013, he was diagnosed with NHO and etidronate was administrated for four months in a previous rehabilitation unit. Serum alkaline phosphatase (sALP) levels

were almost normalized and no further progression of NHO was noted in follow-up radiography (Fig. 1). The administration of etidronate was stopped. However, he continued to experience pain in his left hip joint, so celecoxib was prescribed. At the time of admission to our rehabilitation unit on December 20, 2013, he was suffering from severe chronic pain and LOM in the affected joint without swelling or heatness. Initial sALP levels were 192 IU/L and follow-up sALP level was almost unchanged. The size of NHO had not progressed in radiography. He could not sit upright in a wheelchair due to pain and he could barely sit in a semi-recline position for 10 minutes. The modified Ashworth Scale was grade 0 for all joints except the left hip, which was not assessed due to pain. When pain levels were measured on the visual analogue scale (VAS), the initial VAS was 8 and intermittently 10. Passive range of motion of the left hip was restrictively measured using a goniometer in a supine position due to pain (Table 1). The manual muscle test score was grade III for all extremities.

He could walk using a walker. Gait speed was evaluated using the 6-minute walk test (6MWK) and gait endur-



Fig. 1. (A) Radiograph of the hip showed before extracorporeal shock wave therapy (ESWT). (B) Radiograph of the hip showed after ESWT.

Table 1. Passive ROM of the left hip joint before and after ESWT

Passive ROM of the left hip	Before ESWT (°)	After ESWT (°)	4 weeks after ESWT (°)	12 weeks after ESWT (°)
Flexion	30	70	80	80
Extension	0	0	0	5
Abduction	20	20	20	25
Adduction	20	20	20	20
Internal rotation	50	50	50	50
External rotation	-5	0	0	0

ROM, range of motion; ESWT, extracorporeal shock wave therapy.

ance, which was assessed on a 70-m track in our physical therapy treatment room, was evaluated by the distance that the patient could walk before stopping due to fatigue or pain. The time taken to complete the first lap of this track during the gait endurance test was also recorded. The initial time for the 6MWK was 17 seconds. The distance walked in the gait endurance test was 72 m, with the patient stopping due to hip joint pain. The time taken to walk the first 70 m in the gait endurance test was 272

seconds (Table 1).

Celecoxib was discontinued, and 1 week later ESWT was applied to the anterior aspect of the left hip using Dornier AR2 (Dornier MedTech, Wessling, Germany) (Fig. 2) guided by ultrasound (Fig. 3A). Each application of ESWT involved 4,000 shocks delivered at a rate of 3 Hz and had an energy flux density (EFD) from 0.056 to 0.068 mJ/mm². ESWT was administered once a week for 3 weeks. Physical modalities were not performed.

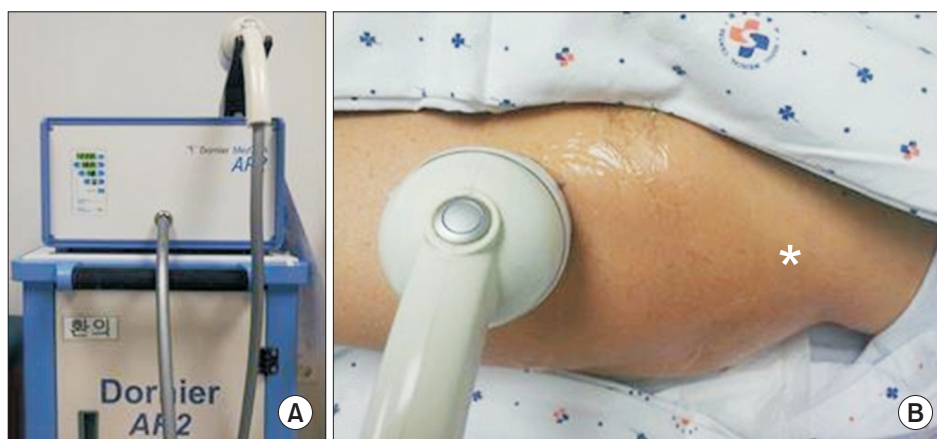


Fig. 2. (A) Dornier AR2 (Dornier MedTech, Wessling, Germany). (B) Extracorporeal shock wave therapy was applied below the anterior superior iliac spine (asterisk).

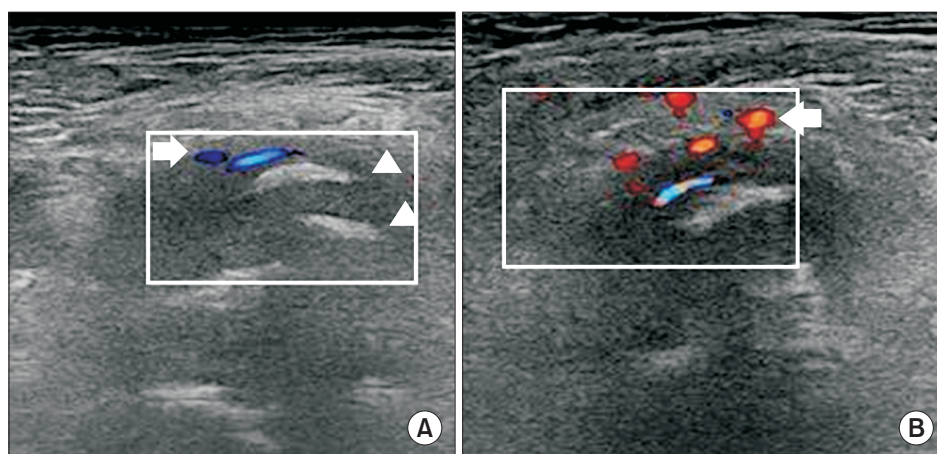


Fig. 3. (A) Transverse ultrasound image taken before extracorporeal shock wave therapy (ESWT) showed multiple hyperechoic foci with no shadowing (arrowhead), vascularity was presented in the color Doppler image (arrow). (B) Blood supply was increased in the color Doppler image (arrow) 4 weeks after ESWT.

Table 2. Measurements of clinical outcome before and after ESWT

Clinical outcome	Before ESWT	After ESWT	4 weeks after ESWT	12 weeks after ESWT
VAS	8-10	3	3	3
6-minute walk test (sec)	17	15	13	14
70 m travel time (sec)	272	195	198	203
Maximum walking distance (m)	72	140	153	203
sALP (IU/L)	192	172	172	186
Sitting time in a wheelchair (min)	10	60	90	120

ESWT, extracorporeal shock wave therapy; VAS, visual analogue scale; sALP, serum alkaline phosphatase.

After ESWT, the passive flexion angle of the left hip joint was 70° (Table 1), the sALP level was 172 IU/L, and the VAS was 3. The duration for which he could sit in a wheelchair was 60 minutes, and he was able to sit comfortably and raise his left foot onto a footrest. The time for the 6MWK was 15 seconds, the distance walked in the gait endurance test was 140 m, and the time taken to walk the first 70 m in the gait endurance test was 195 seconds (Table 2). Radiographs were similar to those obtained pre-ESWT (Fig. 1).

Four weeks after ESWT, sALP levels and VAS were maintained and other outcomes further improved (Tables 1, 2). Color Doppler images obtained 4 weeks after ESWT showed an increase in vascularization around the NHO, but no change in the hyperechoic foci (Fig. 3B). The therapeutic effects were maintained at 12 weeks after ESWT (Table 1). The patient did not request any NSAIDs after ESWT. No complications such as edema, hematoma, dizziness, and paresthesia were noted.

DISCUSSION

The most common clinical findings in NHO are LOM and peri-articular swelling due to interstitial edema of soft tissues around the affected sites [1]. NHO, especially after TBI, is not prone to regression and is hard to treat [8]. The growth of NHO results in the mass-effect, compression of the vessels and nearby peripheral nerves [6]. In this case, NHO was already relatively advanced when etidronate was administered. The NHO was large and would likely have resulted in a mass-effect that was aggravated as time passed. This would have damaged the soft tissues surrounding the NHO through hypoxic injury, inflammatory reactions and tissue necrosis, resulting in pain [7]. The mass-effect likely also triggered sequential inflammatory tissue damage and neurogenic inflammation.

Shockwaves with a low EFD can induce the ingrowth of neo-vessels and tissue proliferation associated with the early release of angiogenesis-related factors including nitric oxide (NO) and vascular endothelial growth factor [3,4]. The neovascularization may lead to improved blood supply and play a role in tissue regeneration. In our case, vascularity around the NHO was increased in the color Doppler images taken 4 weeks after ESWT (Fig. 3B). The local blood supply increased by ESWT around

the NHO may lead to restoration of oxygen, growth factor and nutrients and thus cause the hypoxic tissue to return to aerobic metabolism [4,6,7].

Shockwaves with a low EFD are also able to regulate the inflammatory cascade by modulating NO complex in soft tissues [3]. ESWT may induce a significant increase in NO and a decrease in pro-inflammatory substances, such as nuclear factor kappa B. NO may exert either an excitatory or an inhibitory effect on neuronal transmission [9] and ESWT may reduce the calcitonin gene-related peptide, considered as the main excitatory nociceptive transmitter [5,9]. Also, Wess [10] suggested that ESWT may reorganize pathologic memory giving rise to chronic pain, thus resulting in pain relief. These may explain competent management of chronic painful NHO after ESWT without the diminution in NHO (Fig. 1). Flexion of the left hip joint was particularly improved. We thought the reason was related to the location of NHO. Because NHO occurred in the anterior portion of the left hip joint, more relative limitation of flexion in the left hip joint would occur by mass-effect and the flexion of the left hip joint would significantly improve after ESWT. Also, he had progression in gait speed, gait distance and gait performance. The number of stops caused by pain was reduced during walking. Therefore, the participation in exercise improved and the total amount of exercise were increased.

The study has a number of limitations. An electrodiagnostic study was not performed to confirm nerve injuries around the area of NHO. However, the area of NHO was located a distance from the femoral or the sciatic nerve and no paresthesia was noted along cutaneous branches of their nerves. We did not check the isoenzyme of ALP, but sALP levels were not noticeably different after ESWT. The effects of passive range of motion exercise may have influenced the improvements observed. However, these exercises had been performed since the TBI, which were several months before the ESWT. It is therefore unlikely that the passive range of motion exercises had any effect on the improvements seen after 3 weeks of ESWT. Color Doppler images obtained 4 weeks after ESWT showed increased blood supply. However, power Doppler sonography was not performed and thus it was not possible to quantify the neovascularization. Further quantitative analysis of the neovascularization is therefore required.

In conclusion, this case suggests that ESWT may be

an effective, safe, and easy-to-use treatment for chronic painful NHO after TBI.

CONFLICT OF INTEREST

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

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