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**Resection with limb salvage in an Asian male adolescent with Ewing’s sarcoma: A case report**

Lai CY *et al*. Treatment for adolescent Ewing’s sarcoma

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**Abstract**

BACKGROUND

Ewing’s sarcoma is a highly malignant primary bone tumor that commonly affects children. For young patients, multidisciplinary treatment and limb salvage are recommended, and surgical plans considering the growth potential and bone activity after tumor resection are essential.

CASE SUMMARY

An 11-year-old Asian boy had a 1-mo history of a right-sided limping gait. Imaging revealed a proximal tumor with bone destruction and physeal involvement over the right femoral neck. He was diagnosed with stage IV (T1N0M1aG3) Ewing’s sarcoma with bilateral lung metastases. Neoadjuvant chemotherapy decreased the tumor size and confined it to the metaphyseal region. The patient underwent four stages of surgery: wide tumor excision plus reconstruction with vascular fibular bone graft plus internal fixation; repeat open reduction and internal fixation; femoral lengthening with orthosis after physeal maturity; and orthosis removal and bone elongation (approximately 6 cm). Following surgery, he could walk without discomfort and had almost equal-sized bilateral femoral heads, indicating physis preservation. The surgery was successful, and normal femoral head growth was achieved after complete remission. The patient was able to resume normal activities with equal length of the bilateral lower limbs.

CONCLUSION

Tumor treatment and reconstruction following resection are important in skeletally immature patients with Ewing’s sarcoma to improve quality of life.

**Key Words:** Ewing’s sarcoma; Lower limb discrepancy; Orthosis; Vascular fibular graft; Reconstruction; Case report

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**Core Tip:** Ewing’s sarcoma is an uncommon primary malignant bone tumor, and its treatment is a challenge to the orthopedic surgeon. Beyond survival, current treatment also focuses on functional preservation and cosmetic appearance. This 9-year follow-up case illustrates the complete treatment course from resection to reconstruction and rehabilitation for Ewing’s sarcoma, which demonstrates a complete clinical picture for pediatric specialist and pediatric surgeons dealing with complex tumor surgery. The patient not only achieved complete remission, but also had good functional outcomes with limb salvage.

**INTRODUCTION**

Ewing’s sarcoma is a primary malignant tumor that usually occurs in children. It was first described in 1921 by Ewing J. Ewing’s sarcoma constitutes 17.2% of primary malignant bone tumors in the Caucasian population. In a previous study, the ratio of Ewing’s sarcoma to osteosarcoma was 0.1 in the Chinese group and 0.31 in the American group[1].

A previous study of primary bone cancer in Taiwan conducted between 2003 and 2010 showed that Ewing’s sarcoma accounted for only approximately 8% of all cases[2]. The tumor most commonly occurs in the long bone diaphysis, and the most common site is the femur (22%), followed by the long bones of the lower extremities, including the femur (30%) and the pelvis (19%)[3-5].

Ewing’s sarcoma generally has a favorable prognosis. The Cooperative Ewing’s Sarcoma Study Group reported a 10-year event-free survival rate of approximately 53% in 301 patients with a median age of 15 years. This predilection for the pediatric population and good prognosis underlines the importance of quality of life in addition to survival[6]. However, the Enneking staging system, the most popular surgical staging system for musculoskeletal sarcomas, is not applicable to Ewing’s sarcoma, probably because it does not originate from the mesenchymal connective tissue of the musculoskeletal system, indicating the uniqueness of this tumor[7].

With respect to treatment, a multidisciplinary approach involving neoadjuvant chemotherapy, followed by tumor resection and adjuvant chemotherapy, is required. Non-metastatic, non-recurrent cases have an expected survival rate of 60%-80%[4]. Postoperative radiotherapy has no survival benefit in patients with metastases but improves survival for those with positive surgical margins[8]. Physical function and cosmetic appearance have also become more important in recent years. However, reconstruction after wide resection is difficult in adolescents because of the growth potential and activity.

**CASE PRESENTATION**

***Chief complaints***

An 11-year-old boy presented with a 1-mo history of a right-sided limping gait.

***History of present illness***

The patient fell approximately 1 mo prior to admission. Since this accident, he had persistent pain in his right hip and a right-sided limping gait.

***History of past illness***

He had a known diagnosis of bilateral pes planus.

***Personal and family history***

The patient’s mother was G2P2A0; he was born *via* normal spontaneous delivery at a gestational age of 39 wk. His birth weight was 3300 gm. His parents did not have any specific medical or cancer history.

***Physical examination***

Physical examination revealed swelling of the right proximal femur and hip, tenderness over the right hip and lateral thigh, right hip pain with passive range of motion, and a right-sided limping gait due to weight-bearing pain. He had no loss of muscle power and sensation in the right lower limb.

***Laboratory examinations***

Blood tests revealed the following: lactate dehydrogenase, 401 IU/L; alkaline phosphatase, 158 IU/L; and C-reactive protein, 0.93 mg/dL. Chromosome analysis reported normal male 46,XY karyotype.

***Imaging examinations***

Radiography showed focal bone destruction and osteolytic change over the right femoral neck (Figure 1A). Magnetic resonance imaging showed an intra-skeletal mass in the metaphyseal region with physeal involvement (Figure 1B). Chest computed tomography also indicated bilateral lung metastases (Figure 2).

**MULTIDISCIPLINARY EXPERT CONSULTATION**

We discussed the case with an expert pediatric hematologist (Wu KH, MD, Chief Doctor, Professor, at the China Medical University Hospital) after tumor biopsy reported Ewing’s sarcoma. Neoadjuvant chemotherapy was recommended, and adjuvant chemotherapy was also needed after tumor resection.

**FINAL DIAGNOSIS**

Based on these findings, the patient was diagnosed with stage IV Ewing’s sarcoma (T1N0M1aG3). Histological examination of an incisional biopsy specimen confirmed the diagnosis of Ewing’s sarcoma/primitive neuroectodermal tumor, and the tumor was positive for CD-99 and periodic acid-Schiff staining.

**TREATMENT**

Neoadjuvant chemotherapy with cisplatin, cyclophosphamide, doxorubicin, etoposide, ifosfamide, and vincristine sulfate decreased the tumor size, with the tumor confined to the metaphyseal region (Figure 3). The patient then underwent four stages of surgery.

***Stage 1***

Wide excision of the tumor (physeal preservation) plus reconstruction with a vascular fibular bone graft plus fixation with a locking T-plate (Figure 4). The patient was placed in the supine position, and an incision from the lateral aspect of the thigh was made *via* the standard lateral approach. The tumor was then dissected and subjected to intra-operative frozen section pathology. After complete margin resection was confirmed, the femur and proximal femur were removed to harvest the vascular bone graft from the right fibula *via* the standard lateral approach. The distal ends of the peroneal artery and vein were divided, and the dissection was carried on toward the cephalic end. The vessel branches were ligated, and the flap was harvested based on the right peroneal artery and vein. The total length of the fibula flap was 17 cm, leaving 5 cm distally and 5 cm proximally. The graft was divided into two parts: proximal (10 cm with main blood supply) and distal (6 cm for augmentation of contact with the femoral head). The vascular fibular graft was transferred to the femur, and a vascular anastomosis of the branch of the lateral femoral circumflex artery and vein was performed. Then, the vascular fibular graft was fixed with an Arbeitsgemeinschaft für Osteosynthesefragen (AO) locking plate at the right hip and the distal tibial and fibular with a screw to ensure ankle stability.

The resected mass obtained after performing stage 1 surgery showed a tumor in the metaphysis and adjacent periosteal soft tissue with patchy myelofibrosis and avascular necrosis, but no residual malignant cells, lymphovascular permeation, or perineural invasion. A bone-destructive lesion measuring 3.5 cm × 1.6 cm was found over the distal metaphyseal site of the femur.

After 5 mo, the patient started with gradual limited weight bearing on the right hip. After 14 mo, partial weight bearing (over 30 kg) and walking with a crutch were achieved. However, the plate deformed with broken screws after an increase in partial weight bearing (Figure 5A).

***Stage 2***

After 16 mo, we performed a repeat open reduction and internal fixation for the plate impending failure and malalignment of the vascular fibular bone graft with the AO proximal humeral internal locking system (PHILOS) plate (Figure 5B). The femoral head had enlarged with age, but there was also a limb length discrepancy. After a 7.5-year follow-up period since the tumor excision surgery, a symmetric bilateral hip joint surface and equal femoral head size were noted on radiography. There was also no osteonecrotic change in the right femoral head. The right and left lower limb length were 77 and 83 cm, respectively, with a lower limb discrepancy of 6 cm.

***Stage 3***

Femoral lengthening with orthosis with a distraction rate of 1 mm/day (Figure 5C).

***Stage 4***

Orthosis removal. The AO less-invasive stabilization system locking compression plate elongated the bone by approximately 6 cm.

**OUTCOME AND FOLLOW-UP**

Two months postoperatively, plain film radiography showed complete corticalization of the elongated femur (Figure 5D). The last follow-up plain film was obtained 14 mo postoperatively, and it showed good bone formation of the elongated femur (Figure 5E). The Musculoskeletal Tumor Society score was 28, and the Harris hip score was 96, 16 mo postoperatively[9,10].

**DISCUSSION**

Data on the long-term outcomes of Ewing’s sarcoma are currently lacking[2,11]. In Taiwan, Ewing’s sarcoma accounted for only 8% of all bone cancer cases during 2003-2010[12]. We reported the case of an adolescent patient with Ewing’s sarcoma who underwent three types of surgery. The first was a successful hip joint preservation, with the final imaging during follow-up showing an equal femoral head diameter and joint surface congruity. The second was graft substitution of the femoral neck with vascular autograft for the preservation of growth potential, and the choice of bone and implant for augmentation and fixation. The third was femoral lengthening technique selected to enable the patient to regain normal ambulatory capacity.

Despite novel treatment modalities, oncological surgery remains crucial for improving survival. In addition, both patients and their family are concerned about regaining normal function and cosmetic appearance after tumor resection. Given that Ewing’s sarcoma affects the bones and soft tissues, potential functional changes are an important consideration when planning surgery. The three main growth areas in the proximal femur are the physeal plate, the growth plate of the greater trochanter, and the femoral neck isthmus. The proximal femoral physeal plate contributes to approximately 30% of the overall length of the femur, and 13% of the entire lower limb[13]. It is important to preserve joint function and activity if limb preservation can be expected, especially in pediatric patients who still have growth potential.

A malignant tumor in the proximal femur can affect the hip joint. Once we decided to perform limb salvage surgery, age, growth potential, and physis involvement were the first factors to be considered. Tsuchiya *et al*[14] used a vascular fibular graft to allow growth of the physis in the fibula in the hip joint, with cadaveric femoral allograft as reinforcement. However, this method has not yet been proven to effectively enable growth. Intra-epiphyseal excision is sometimes an option for long bone tumors[15]. In our case, we selected a hip joint preservation technique. Tumor resection in the proximal femur, particularly the femoral neck, arthroplasty, or vascular bone graft with allograft bone augmentation as a structural graft, can be performed in adult patients[14,16].

Although vascular grafts allow for bony growth in pediatric patients, surgery allows enlargement of the hip joint; thus, hip-preservation surgery is needed. Furthermore, the small diameter of the femoral neck makes it challenging to obtain an appropriate proximal femoral allograft in this patient population. Re-implantation of the resected proximal femur bone post cryotherapy is another option to reinforce the biomechanical construct[13]. We used a folded autologous vascular fibula graft reconstructed from the proximal femur, particularly the femoral neck. However, given that the tumor was not resected from the femoral subtrochanteric region and that the patient was only 11 years old, we used a locking plate to stabilize the vascular fibula graft[17]. The Capanna technique was not suitable in our case, because the literature on pediatric allograft is limited, and the smaller bone diameter makes this technique challenging, as it involves a sandwich-like bone graft. If we performed a reconstruction technique similar to that for adults, hemi-arthroplasty is an option with a low risk of mechanical failure. However, the bony growth of the proximal femur in our patient did not reach the expected length, and thus, the implant failed. Accordingly, we performed another surgery with AO PHILOS after 1.5 years. This allowed for better biomechanical distribution of the head-neck force to the femoral shaft, and the vascular graft successfully combined.

After 8 years, the diameter of the femoral head was equal to that of the other site, indicating that preservation of the physis allowed for enlargement of the hip joint. However, the length discrepancies remained an issue. Similar to those in the tibia, complications, particularly infection, in extra-skeletal fixation are of great concern[18,19]. Concerning the femoral lengthening device, an extra-skeletal mono-lateral system, such as orthosis, is an efficient approach that does not require joint involvement. The system avoids broken screws and allows for preservation of the remaining physis[20]. One study compared limb length discrepancies among elongation devices in 73 patients with Ewing’s sarcoma or osteosarcoma, and found that 10 patients required secondary lengthening. The average length achieved was 8.1 cm, and the final residual discrepancy was 1.5-2.5 cm.

Angulation, torsion, osteoporosis, joint instability, muscular weakness, and patient and family cooperation are factors that must be considered prior to long bone elongation surgery. In general, the patient needs to undergo the following five phases before removing the orthosis: (1) Osteotomy; (2) Latency; (3) Distraction; (4) Neutralization; and (5) Consolidation[19]. Physical activity is limited during the consolidation phase and the percutaneous pins of orthosis also increase the risk of pin tract infection. An internal fixation with a locking plate prevents these problems. The patient underwent another surgery after the complete distraction phase to change the orthosis to an internal fixator during the consolidation phase.

**CONCLUSION**

Each surgical modality used to treat Ewing’s sarcoma has its own challenges and risks of failure or complications. In the current case, tumor resection, reconstruction, rehabilitation, elongation, and rehabilitation within 9 years enabled good recovery of the patient. Corticalization was confirmed on the final radiographic follow-up, and the patient is currently doing well, with good physical function.

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**Footnotes**

**Informed consent statement:** The patient and his family agreed to the publication of this case report and provided a signed informed consent form.

**Conflict-of-interest statement:** All the authors report no relevant conflicts of interest for this article.

**CARE Checklist (2016) statement:** The authors have read the CARE Checklist (2016), and the manuscript was prepared and revised according to the CARE Checklist (2016).

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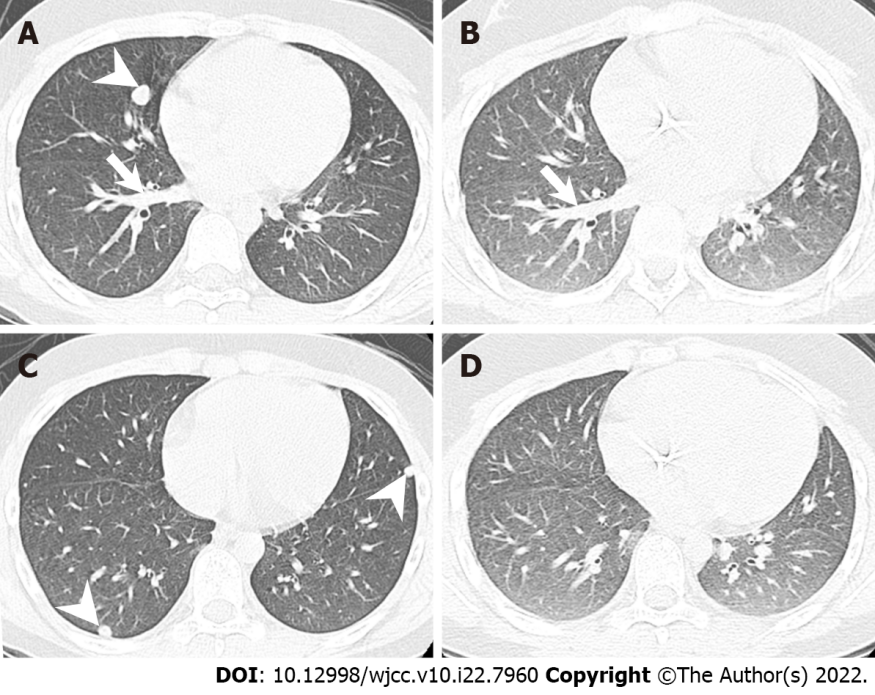
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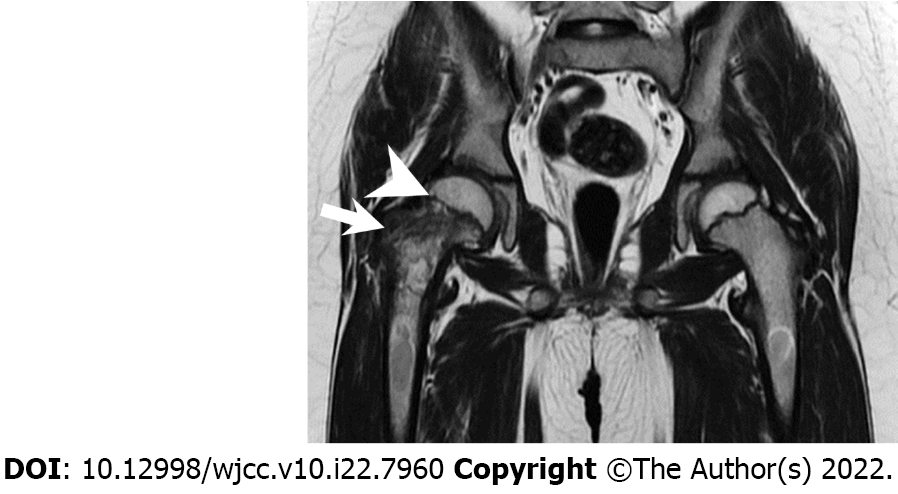
**Figure Legends**



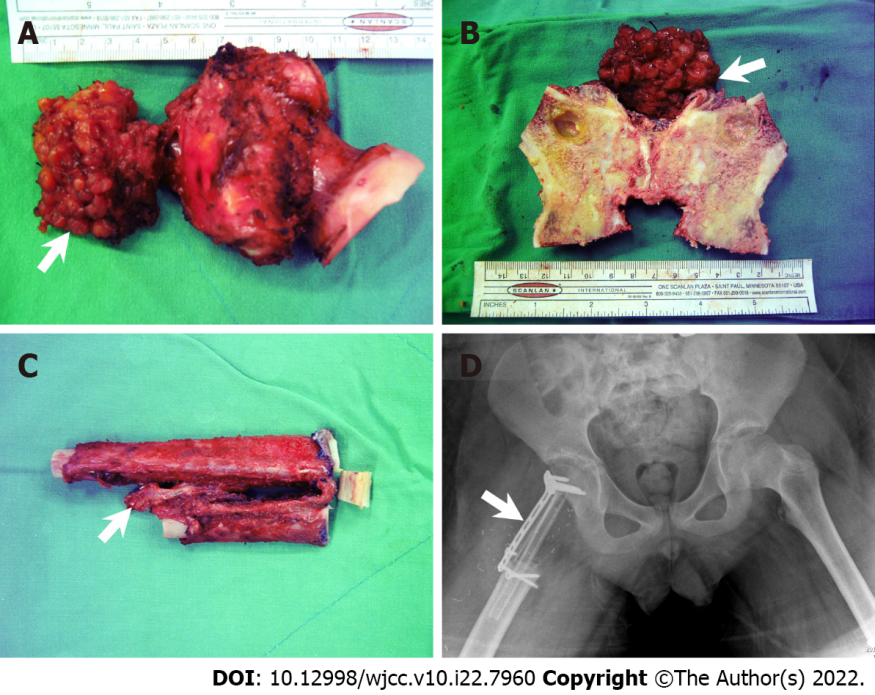
**Figure 1** **Radiography and magnetic resonance imaging.** A: Initial plain X-ray film of the pelvis (anteroposterior view). The arrow indicates the location of the tumor with cortical reaction at the femoral neck region; B: Initial pelvis coronary T2 STIR image showing involvement of the physis. The arrow and arrowhead indicate the location of the tumor and the femoral physis, respectively.

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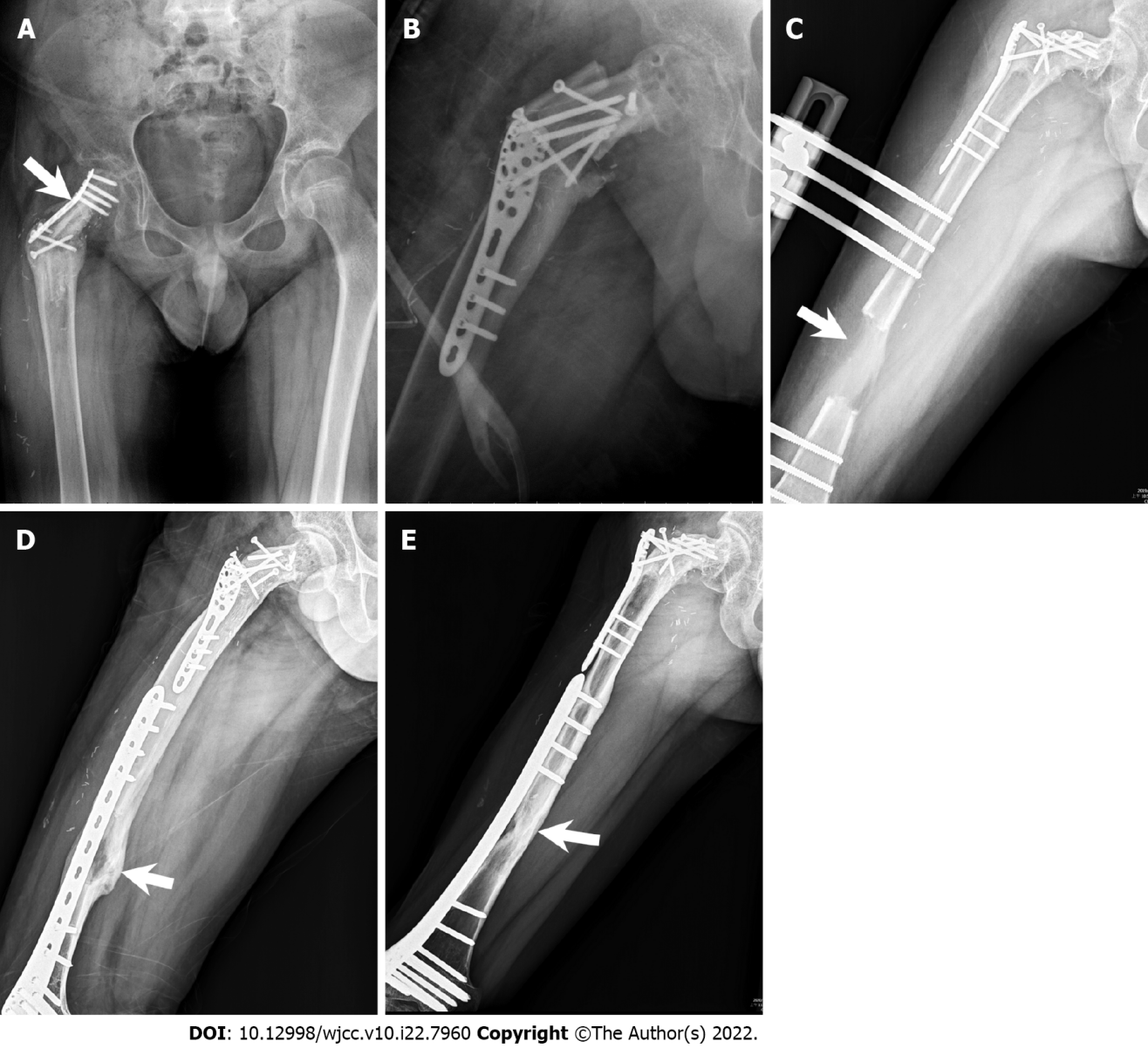
**Figure 2 Initial chest computed tomography and chest computed tomography after neoadjuvant chemotherapy.** Full remission of the lung metastases after the neoadjuvant chemotherapy is noted. A: Initial: The metastatic Ewing’s sarcoma over the right middle lobe (arrowhead). Pulmonary artery is not distended (arrow), which implies no obstructive tumor thrombus in this multiple metastasis situation; B: After neoadjuvant chemotherapy: Localized same level chest computed tomography with the pulmonary artery (arrow). The previous tumor had disappeared; C: Initial: The metastatic Ewing’s sarcoma in the right inferior lobe (arrowhead) and left inferior lobe with another metastatic Ewing’s sarcoma (arrowhead); D: After neoadjuvant chemotherapy: Localized same level chest computed tomography with similar pulmonary artery distribution. The previous tumors had disappeared.

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**Figure 3 T2 magnetic resonance image (coronary view) after neoadjuvant chemotherapy.** The arrow and arrowhead indicate the location of the tumor and the femoral physis, respectively. Note that the tumor is now confined only to the physis.

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**Figure 4 Intraoperative sample and postoperative pelvic plain imaging.** A: Gross view of the resected tumor and resected femoral neck (arrow: Ewing’s sarcoma extra-bony part); B: Split resected femoral neck; C: Folded autologous fibular graft (arrow: vascular pedicle); D: Pelvic plain imaging (anteroposterior) obtained immediately after the surgery (arrow: locking plate used to fix the fibular graft).

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**Figure 5** **Radiographic images.** A: Plain X-ray film obtained after an increase in partial weight bearing. The plate was deformed (arrow: bending site of the plate); B: Revision and fixation with Arbeitsgemeinschaft für Osteosynthesefragen proximal humeral internal locking system plate; C: Radiographic image of orthosis with complete distraction. The femoral distraction gap is approximately 6.5 cm (arrow: distraction gap); D: Radiographic image showing removal of the orthosis and international fixation with less-invasive stabilization system complete corticalization (arrow: corticalized bone); E: Radiographic image showing complete consolidation phase (arrow: cordialized bone with increased radiopacity).



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