Renal cell carcinoma (RCC) is the most common malignancy of kidneys in adults and accounts for 2-3% of all cancers worldwide [1]. Almost one-third of the patients present with metastatic disease and another 20-30% develop recurrence or metastasis after nephrectomy [2]. Majority of metastasis present in the first few years after the treatment of RCC. Late metastasis and recurrences after 10 years of curative treatment is a known biological behavior of RCC and a long follow-up is required for the detection of metastasis. Late solitary bony metastasis is very rare. No case of a late solitary metastasis of distal femur, treated with wide local excision and reconstruction, is available in the literature. We present a case of solitary metastatic lesion of distal femur 12 years after radical nephrectomy for renal cell carcinoma in a 64-year-old male.

Case Report: The patient presented to us with swelling in the right distal thigh for three years with a history of radical nephrectomy for RCC 12-years back. The lesion was not responding to local radiotherapy, and chemotherapy in the form of oral pazopanib, taken before orthopedic consultation. After core biopsy, the tumor was managed by intercalary wide local excision and reconstruction using a cement block and a lateral locking plate. Post-operatively, the histopathology report confirmed the diagnosis to be a clear cell tumor, consistent with metastatic RCC. The patient is independently mobile and tumor-free 2 years after the surgery.

Conclusion: Wide resection and reconstruction of the skeletal defect remains the mainstay of the management of metastatic solitary lesion. We have presented a unique case of distal femoral solitary metastatic deposit from a primary RCC 12 years post radical nephrectomy treated by intercalary resection and reconstruction with bone cement-plate hybrid construct.

Keywords: Renal cell carcinoma, metastasis, femur, wide local excision, intercalary resection.
using intercalary resection and knee joint preservation with a cement block augmented with a plate and screw construct was carried out.

**Case Report**

A 64-year-old male presented to us with the complaint of swelling in the right distal thigh for the last three years. It was gradually increasing in size with occasional dull aching pain. He had a right-sided radical nephrectomy for RCC 12-years back. On examination, the swelling was in the anteromedial aspect of right distal thigh, firm to bony hard in consistency, globular, immobile and deep to the muscle plane. It measured 7 × 10 cm and the range of motion of the knee joint was unaffected.

The pre-operative radiograph showed an eccentric lytic lesion in the distal femur with moth-eaten appearance and involvement of surrounding soft tissue (Fig. 1). Magnetic resonance imaging revealed a well-circumscribed mass in the distal femur, 11 × 4 × 5 cm in dimensions, iso-intense on T1 weighted image and hyper-intense on T2 weighted image. It was located 25 cm from the tip of the greater trochanter and 5 cm proximal to articular surface laterally and 3 cm medially (Fig. 2).

PET scan showed no other lesions in the body (Fig. 3). There was no vascular encasement by the tumor on CT angiogram. Core biopsy confirmed the diagnosis of metastatic clear cell RCC. He had received local radiotherapy 1-year back and was started on oral Pazopanib before orthopedic consultation was sought, but the tumor showed no regression in its size. Pre-operative planning for resection of the tumor with the reconstruction of the intercalary defect with bone cement was done and exact margins of resection were decided (Fig. 4). Exposure was achieved by anterior midline approach and a plane was developed between the tumor and the vastus intermedius muscle. Wide local excision of the tumor with its intact capsule, along with the cuff of surrounding vastus intermedius muscle was carried out (Fig. 5). There was no breach of the tumor capsule during excision. The intercalary defect of 16 cms was filled up with Simplex P bone cement (Stryker, USA) after keeping two intra-medullary 4.5 mm Steinmann pin medially. Laterally, it was supported by a 13-hole titanium locking plate before cement was set. Post-operative plain radiograph was done (Fig. 1). The excised segment of bone including the muscle cover measured about 16.5 × 7 × 5 cm.
RCC responds poorly to radiotherapy, chemotherapy, and immunotherapy. These modalities have been used in the treatment of metastatic RCC but the response rate is poor making surgical excision treatment of choice wherever possible [7]. Surgical excision of a solitary metastasis in RCC may provide five-year survival rates of 35-60% [7]. Factors improving survival after excision of bone metastasis are a solitary metastatic disease, younger age at presentation, tumor-free resection margins, absence of a pathological fracture, histology grade, and longer time from nephrectomy to bone metastases [8]. Fuchs et al. [7] studied 60 patients with solitary metastatic RCC of which 13 patients were managed by wide.

Intra-operative frozen section samples from bony and soft tissue margins were clear of any tumor cells. Post-operatively, the histopathology report confirmed the diagnosis to be metastatic clear cell tumor of the right distal femur, consistent with metastatic RCC (Fig. 6). The patient was allowed weight-bearing as tolerated from the first post-operative day and knee range of motion exercises were started within limits of pain. At 2 years follow-up the patient walks without support, with knee flexion up to 120 degrees without any complaints or complications (Fig. 7). A follow-up PET scan did not show any active lesion in the body (Fig. 8).

Discussion

These late metastases present a unique diagnostic dilemma, and a primary bone tumor or metastasis from any other site, has to be ruled out at this age. This includes a planned biopsy from the lesion. Our patient was treated with intercalary excision, because it was located away from the knee joint and thus, salvage of the native knee joint was possible. Reconstruction options with salvage of knee joint included bone transport, allograft reconstruction, bone grafting, or custom-made metallic spacer. Primary cement spacers for intercalary reconstruction have been described by Puri et al. [6]. This proved early mobility to the patient which was crucial in a patient of this age. The rehabilitation period was very short as this patient was allowed weight bearing within limits of pain within 24 h after the surgery and has been walking independently since then. The cost of the treatment is also lesser than the other mentioned treatments. Allografts will carry a risk of resorption, infection or fracture. Bone transport takes a longer period of time and may have failed considering that this patient was elderly, was on immunotherapy, and had received chemotherapy and local radiotherapy in the past. These spacers can be easily revised to a vascularized fibular graft or a custom-made intercalary prosthesis in the future if needed.

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Pazopanib is an oral angiogenesis inhibitor acting on vascular endothelial growth factor receptor, platelet-derived growth factor receptor and tyrosine kinase receptor. It has shown to improve progression-free survival [9]. However, our case did not show any good clinical response with pazopanib in terms of containment and regression of the lesion.

Bony metastasis is relatively common in primary metastatic RCC and may be seen in 35% of these patients [10]. Swanson et al. studied 947 patients with RCC, and found skeletal metastasis in 26.7% of patients, mostly involving spine, pelvis and proximal femur [10]. It is generally associated with widespread metastasis and isolated skeletal metastasis is uncommon [9]. It is also very rarely seen as an isolated late metastasis. Miyao et al. [3] studied the pattern of delayed metastasis in 470 patients who had undergone curative treatment of RCC. In this series, 30 patients (6.4%) developed a late metastasis. Only six sites of bony involvement were found of which only one was treated surgically. A case of metastatic lesion of tibia and ankle from primary RCC 1-year post-nephrectomy managed surgically by above-knee amputation has been described before [11]. Late metastasis of an RCC in soft tissues of the thigh has also been reported [12]. Rare cases of skeletal muscle metastasis from RCC have also been reported [13, 14, 15, 16].

After a thorough literature search, distal femoral solitary metastatic deposit from primary RCC 12 years post radical nephrectomy remains a rare entity to the best of our knowledge. Thus, the uniqueness of our case lies in firstly the site being distal femur around 4.9 cm above the articular surface, secondly, the time of presentation which is 12 years post nephrectomy for primary RCC. Moreover, the technique employed for the surgical management of the tumor by resection and reconstruction of the intercalary defect with cement block is also quite uncommon. Although there were no signs of loosening of the cement spacer in this case, one should be careful to look for radiolucency around the cement bone junction and pain while following up these cases. The choice of this reconstruction technique is justified by the benefit of immediate stability, early mobilization and cost-effectiveness. Moreover, looking at the poor prognosis and life expectancy of metastatic RCC this reconstruction technique may be a viable option. The post-operative histopathological diagnosis was metastatic clear cell variety of RCC. The histological section revealed infiltrative tumor cells, arranged in diffuse large sheets showing scattered pleomorphic large cells with abundant clear cytoplasm, prominent nucleoli, and round nucleus. Atypical mitotic figures indicating a highly proliferative tumor with prominent multiple small-caliber blood vessels and diffuse areas of necrosis with hemorrhages were also well marked (Fig. 6).

**Conclusion**

Usually, patients with metastatic solitary lesion from primary RCC have longer survival, hence wide local resection and reconstruction of the skeletal defect remains the mainstay of management to prevent local disease progression and further complications. Intercalary re-section and reconstruction with bone cement-plate hybrid construct is an easy, safe and economical method of reconstruction for metaphyso-diaphyseal tumors where joint salvage is the aim.

### Clinical Message

Solitary bony metastatic deposit of Renal Cell carcinoma, 12 years after radical nephrectomy remains a rare entity. Wide local resection remains the mainstay of management, improving the long-term survival. Reconstruction of the intercalary bone defects with bone cement-plate hybrid construct remains a viable option in cases where the life expectancy is relatively short or when the patient cannot afford a prosthesis construct. Bone cement-plate hybrid construct restores the limb function and improves the quality of living.
Conflict of Interest: Nil
Source of Support: Nil

Consent: The authors confirm that informed consent was obtained from the patient for publication of this case report

How to Cite this Article
Choudhury AK, Verma R, Dhyani A, Paul S, Goyal T. Late Metastasis of Renal Cell Carcinoma in Distal Femur Treated with Knee Joint Preservation: A Case Report. Journal of Orthopaedic Case Reports 2022 January;12(1): 1-5.