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A Case Report of Abnormal Fracture Healing as Detected With High-Resolution Peripheral Quantitative Computed Tomography

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Introduction

Fracture healing is a complex repair process with the primary objective of restoring the mechanical function of the fractured bone (1). Quantifying outcome in clinical fracture healing trials remains challenging (2,3). Using high-resolution peripheral quantitative computed tomography (HR-pQCT) in combination with micro finite element analysis (μFEA), we previously described the typical healing of a distal radius fracture. This process consists of an increase in bone density of the trabecular compartment peaking at 6 wk post fracture, corresponding to the formation of a mineralized fracture callus (4,5). In the present case report, we describe a patient who deviated from this usually observed pattern of fracture healing.

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dimethyl sulfoxide 50%, pregabalin, calcitonin, paracetamol, and tramadol. At the last follow-up, 3 yr post fracture, active motion of the left wrist was 0° in all axes and both spontaneous pain and allodynia were present, although the patient was able to use her affected hand to a greater extent during normal daily activities than before treatment.

Routine screening on osteoporosis with dual-energy X-ray absorptiometry and laboratory examination showed osteopenia at the lumbar spine and proximal femur, and secondary hyperparathyroidism (parathyroid hormone 8.0 pmol/L, ref. 1.3–6.8), due to vitamin D deficiency (25(OH) vitamin D: 33 nmol/L, ref. >75 nmol/L). With cholecalciferol supplementation, the latter was resolved by 12 wk post fracture (25(OH) vitamin D: 67 nmol/L, parathyroid hormone 4.5 pmol/L).

**HR-pQCT Measurements**

HR-pQCT is a low-dose radiographic imaging modality with an isotropic voxel size of 82 (XtremeCT-1; Scanco Medical AG, Brüttisellen, Switzerland). The high resolution enables the assessment of bone microarchitecture in vivo (7) and estimation of bone strength μFEA (8).

HR-pQCT scans (XtremeCT-1) were performed at 1, 3, 6, 12, and 115 wk post fracture using the manufacturer’s clinical in vivo settings (effective energy of 60 kVp, tube current of 900 µA, and 100-ms integration time), in accordance with the approved study protocol (NTR3821). From these images, bone density, geometry, and microarchitectural and μFEA parameters were derived (4,5).

**Results**

Incongruent with the healing pattern observed with HR-pQCT described earlier (4,5), a decrease in trabecular density was detected at 6 wk post fracture (−11.9 mgHA/cm³), where the typical healing response showed an increase (median +35.4 mgHA/cm³). This deviation was not restored even after 115 wk (Fig. 1A). Simultaneously, the

![Fig. 1. High-resolution peripheral quantitative computed tomography-derived bone parameters of the presented case of abnormal fracture healing (black circles) compared to the other subjects of the study cohort (white squares, N = 14). In the case patient, the trabecular density is seen to decrease early in the healing process, persisting up to 115 wk (A), along with the trabecular number (B). In contrast to trabecular density, cortical density during fracture healing follows a normal pattern (C), which explains the restored compression stiffness (D). Data are presented as median with interquartile range. Scans were performed (on average) at 1, 3, 6, 12, and 115 wk post fracture.](image-url)
trabecular number declined significantly more from 6 wk post fracture onward (−0.7 vs median −0.2 mm⁻¹), a difference persisting up to 115 wk (−1.5 vs median −0.5 mm⁻¹, Fig. 1B).

In contrast, the cortical region healed normally, following a similar pattern as the other patients in the study (Fig. 1C). As a result, calculated compression stiffness using μFEA was comparable to the other patients (Fig. 1D).

Visual inspection of the HR-pQCT reconstructions confirmed cortical healing. The axial and coronal reconstructions show that the trabecular bone loss in the deviating patient (case) was more extensive than the generally observed decline in trabecular density and structure (Fig. 2 and Supplemental Video S1). This is especially evident in the segmented 3-dimensional reconstructions. Remarkably, these images also indicate that the trabecular bone loss occurred predominantly in the region proximal to the fracture line (Fig. 2).

**Discussion**

The present case report demonstrates the potential of HR-pQCT to distinguish different patterns of fracture healing in a clinical setting. In a single patient, we detected a remarkable resorption of trabecular bone proximal to the fracture line, whereas the cortical bone healed normally. These regional discrepancies were not detected to this extent in the other patients with a fracture of the distal radius.

The relevance of CRPS in the present case report is uncertain. Although regional osteoporosis was part of the CRPS spectrum when it was first described by Paul Sudeck in 1900 (9), current clinical imaging techniques are unable to discriminate normal posttraumatic bone loss as a result of immobilization and disuse from bone loss concurring with CRPS development (10). However, the data presented here are in accordance with the recent suggestion that HR-pQCT could be a solution to this challenge (11): the trabecular bone loss discriminated the patient with CRPS from the other study participants (4) as early as 6 wk post fracture, a period characterized by cast immobilization for all subjects. Nonetheless, CRPS remains an elusive clinical syndrome subject to much debate (12–14). Using HR-pQCT, the incidence and role of localized bone loss within the CRPS population could be further elucidated.

A limitation of the present case report is that it only presents a single affected patient to illustrate the detection of abnormal fracture healing using HR-pQCT. Additionally, based on the current data available to us, the etiology of the observed bone loss is not certain and can only be speculated on. However, this case report indicates that HR-pQCT offers a new way of looking at abnormal fracture healing even before the presentation of clinical symptoms.

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Appendix
Supplementary material

Supplementary data to this article can be found online at doi:10.1016/j.jocd.2017.05.004.

References