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EDITORIAL

The forgotten phase of fracture healing: the need to predict nonunion

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A. H. R. W. Simpson Trauma care has improved dramatically over the last three decades with decreased mortality and improved techniques for soft tissue cover and fracture stabilisation. After trauma, it is recognised that there is a golden hour in which optimum resuscitation is essential. Following the golden hour there is an early phase in which debridement of open wounds and soft tissue cover should be obtained along with fracture stabilisation, usually within 48 hours.¹ Recovery of muscle strength and function starts once the fracture has been stabilised but progresses more rapidly in the rehabilitation phase after fracture union. However, in between the early treatment phase and the rehabilitation phase, there is a prolonged "forgotten" phase during which patients are monitored infrequently except for cast treated fractures having alignment checks² and the patient and clinicians wait for the fracture to heal.

However, at present clinicians treating fractures are powerless in this phase as they do not have a technique for monitoring the early rate of healing. This is compounded by the fact that healing times are known to vary according to the bone, type of fracture and location within the bone.3 For clavicle fractures, symptoms and smoking status can indicate that a patient is at a greater risk of impaired healing, but does not identify individual nonunion.⁴ Unfortunately, radiographs in adults (even when they are used in a standardised fashion)⁵ and even CT scans, typically do not show evidence of fracture union for ten or more weeks. Thus assessing the rate of healing is especially difficult in the first two to three months post-fracture.

Over the past few years, a number of prospective trials have examined the best way to treat various fractures.⁶⁻¹¹ However, even when fracture repair proceeds uneventfully, the considerable morbidity the

patient experiences may be underestimated and it is often several years before full function has returned.¹² If fracture repair does not progress smoothly and a nonunion (or even delayed union) develops, the morbidity is substantially greater and often associated with severe financial hardship for the patient and a large burden for the healthcare system.¹³ Nonunions are often multifactorial¹⁴ and their rates vary but are typically 5% for fractures of the clavicle and tibia, but rise to nearly 10% of fractures in working age adults.¹⁵ The treatment of established nonunions is often complex both surgically¹⁶⁻¹⁹ and biologically.^{20,21} Preventing patients getting to this advanced state would therefore be attractive both from the patient and society's perspective. Yet, the FDA definition of nonunion being a fracture that is ununited at nine months,²² subjects patients to prolonged suffering and waiting until nine months to diagnose a nonunion should be considered a failure of modern fracture treatment. Other definitions include a failure to heal within the expected time and a lack of progression of fracture healing on sequential radiographs, but as this again relies on a radiographic technique, it only brings the time to diagnosis of nonunion down by a few months.

The current long period before we diagnose nonunion is a consequence of our inability to monitor healing in the first few months in this forgotten phase. There is therefore a desperate need for tools, such as that proposed by Kienast et al,23 that can determine in the first few months if a fracture is progressing to a nonunion. If such tools were available, it would transform, the care of fracture patients and the first two months would become a vital stage of assessment and no longer the forgotten phase of fracture repair.

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References

- Lewis PM, Waddell JP. When is the ideal time to operate on a patient with a fracture of the hip?: a review of the available literature. *Bone Joint J* 2016;98-B:1573-1581.
- Mirghasemi SA, Rashidinia S, Sadeghi MS, Talebizadeh M, Rahimi N. A prospective study of a modified pin-in-plaster technique for treatment of fractures of the distal radius. *Bone Joint Res* 2015;4:176–180.
- Sandberg OH, Aspenberg P. Glucocorticoids inhibit shaft fracture healing but not metaphyseal bone regeneration under stable mechanical conditions. *Bone Joint Res* 2015;4:170–175.
- Clement ND, Goudie EB, Brooksbank AJ, Chesser TJ, Robinson CM. Smoking status and the Disabilities of the Arm Shoulder and Hand score are early predictors of symptomatic nonunion of displaced midshaft fractures of the clavicle. *Bone Joint J* 2016;98-B:125-30.
- Leow JM, Clement ND, Tawonsawatruk T, Simpson CJ, Simpson AHRW. The radiographic union scale in tibial (RUST) fractures: Reliability of the outcome measure at an independent centre. *Bone Joint Res* 2016;5:116–121.
- Sims AL, Parsons N, Achten XL, et al. The World Hip Trauma Evaluation Study 3: Hemiarthroplasty Evaluation by Multicentre Investigation – WHITE 3: HEMI – An Abridged Protocol. *Bone Joint Res* 2016;5:18–25.
- 7. Huxley C, Achten J, Costa ML, Griffiths F, Griffin XL. A process evaluation of the WHITE Two trial comparing total hip arthroplasty with and without dual mobility component in the treatment of displaced intracapsular fractures of the proximal femur: Can a trial investigating total hip arthroplasty for hip fracture be delivered in the NHS? *Bone Joint Res* 2016;5:444–452.
- 8. Wordsworth M, Lawton G, Nathwani D, et al. Improving the care of patients with severe open fractures of the tibia. *Bone Joint J* 2016;98-B:420-424.
- White TO, Bugler KE, Appleton P, et al. A prospective randomised controlled trial of the fibular nail versus standard open reduction and internal fixation for fixation of ankle fractures in elderly patients. *Bone Joint J* 2016;98-B1248-1252.
- Ersen A, Atalar AC, Birisik F, Saglam Y, Demirhan M. Comparison of simple arm sling and figure of eight clavicular bandage for midshaft clavicular fractures. *Bone Joint J* 2015;97-B:1562-1565.
- Parker MJ. Hemiarthroplasty versus internal fixation for displaced intracapsular fractures of the hip in elderly men. *Bone Joint J* 2015;97-B:992-996.
- Gaston P, Will E, McQueen M, Elton R, Court-Brown C. Analysis of muscle function in the lower limb after fracture of the diaphysis of the tibia in adults. *The Journal of Bone and Joint Surgery [Br]* 2000;82-B:326-331.
- Hak DJ, Fitzpatrick D, Bishop JA, et al. Delayed union and nonunions: epidemiology, clinical issues, and financial aspects. *Injury* 2014;45 Suppl 2:S3-7.

- Mills L, Tsang J, Hopper G, Keenan G, Simpson AHRW. The multifactorial aetiology of fracture nonunion and the importance of searching for latent infection. *Bone Joint Res* 2016;5:512–519.
- Mills LA, Aitken SA, Simpson AHRW. The risk of nonunion per fracture: current myths and revised figures from a population of over 4 million adults. Acta Orthopaedica 2017;88:434–439.
- Gruber HE, Ode G, Hoelscher G, et al. Osteogenic, stem cell and molecular characterisation of the human induced membrane from extremity bone defects. *Bone Joint Res* 2016;5:106–115.
- Wang X, Luo F, Huang Z, Xie Z. Induced membrane technique for the treatment of bone defects due to post-traumatic osteomyelitis. *Bone Joint Res* 2016;5:101–105.
- Burghardt RD, Manzotti A, Bhave A, Paley D, Herzenberg JE. Tibial lengthening over intramedullary nails: A matched case comparison with Ilizarov tibial lengthening. *Bone Joint Res* 2016;5:1–10.
- Rohilla R, Wadhwani J, Devgan A, Singh R, Khanna M. Prospective randomised comparison of ring *versus* rail fixator in infected gap nonunion of tibia treated with distraction osteogenesis. *Bone Joint J* 2016;98-B:1399-1405.
- Ismail HD, Phedy P, Kholinne E, et al. Mesenchymal stem cell implantation in atrophic nonunion of the long bones: A translational study. *Bone Joint Res* 2016;5:287–293.
- Papanagiotou M, Dailiana ZH, Karachalios T, et al. RhBMP-7 for the treatment of nonunion of fractures of long bones. *Bone Joint J* 2015;97-B:997-1003.
- 22. No authors listed. United States Food and Drug Administration Guidance document for the preparation of investigational device exemptions and pre-market approval applications for bone growth stimulator devices. https://www.fda.gov/ohrms/dockets/98fr/98d0238.pdf (date last accessed 13 October 2017).
- Kienast B, Kowald B, Seide K, et al. An electronically instrumented internal fixator for the assessment of bone healing. *Bone Joint Res* 2016;5:191–197.

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A. H. R. W. Simpson: Writing the paper

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