Dupuytren's disease shows profound hematopoietic and mesenchymal stem cell abnormalities involving palmar fat and skin in addition to fascia: Novel implications for pathogenesis and therapy

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Abstract

Background: DD is a common fibroproliferative disease affecting the hand. Myofibroblasts found in the DD palmar fascia are responsible for digital contracture. However, their exact source is unknown. Palmar skin and fat- derived cells as a potential origin of abnormal cells remains underexplored.

Objective: To characterise hematopoietic and mesenchymal stem cells in Dupuytren's disease (DD) fascia, fat and skin compared to carpal tunnel control.

Methods: Nine DD patients with digital fixed flexion deformity were enrolled, and biopsies taken from the diseased cord, nodule, peri-nodular fat, distant palmar fat and the skin overlying the nodule. Fluorescence Activated Cell Sorting (FACS), immunohistochemistry and quantitative real time polymerase chain reaction (QRT-PCR) were used to identify expression of five mesenchymal (CD's 13, 29, 44, 90, 166) and two haematopoietic (CD's 34,117) stem cell markers.

Results: There was a significantly higher expression of CD13 in all examined DD tissue sites compared to controls (p=0.02). In addition CD44 was significantly over expressed in the cord and nodule (p=0.02), while CD34 was over expressed in the skin overlying DD nodules (p=0.008). The mean number of positive cells expressing all stem cell markers were significantly greater in the DD cord compared to carpal tunnel fascia (p=0.003).

Conclusions: This study provides the first detailed characterization of mesenchymal stem cells in DD which are predominantly located in the cord. DD appears to represent a benign

tumour that may potentially result from abnormal mesenchymal progenitor cell expansion than a primary inflammatory disorder. If confirmed, adipodermofasciectomy may be a potentially important primary therapy.