

# Native and Activated Hepatic Stellate Cells Stimulate Liver Regeneration In Rats After Partial Hepatectomy and 2-Acetylaminofluorene Injection

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**Abstract** One the current challenges of modern hepatology is to find new approaches to stimulate liver regeneration and to find new methods for liver diseases treatment. Cell therapies, which are based on using regional stem cells for disease treatment, are under active development. However, studies, devoted to their transplantation, are currently scarce. In recent years hepatic stellate cells are considered to be hepatic stem cells. It is known that activated hepatic stellate cells can transdifferentiate into myofibroblasts and lead to liver fibrosis. The aim of our work was to study the influence of native and activated hepatic stellate cells *in vivo* by lead nitrate injection after transplantation into partial hepatectomized rats, which is considered to be a classical model to study liver regeneration. Injection of 2-acetylaminofluorene (AAF), that selectively eliminates hepatocytes proliferation,

was used to understand the hepatic stellate cells role in liver regeneration process better.

Our results suggest that transplanted native and activated hepatic stellate cells can differentiate into hepatocyte-like cells and positively influence liver regeneration without inducing liver fibrosis.

**Key words** hepatic stellate cells, liver regeneration, partial hepatectomy, 2-acetylaminofluorene

## References

1. Ramadori G (1991) The stellate cell (Ito-cell, fat-storing cell, lipocyte, perisinusoidal cell) of the liver. New insights into pathophysiology of an intriguing cell. *Virchows Arch B Cell Pathol Incl Mol Pathol* 61(3): 147-158.
2. Kordes C, Sawitza I, Müller-Marbach A et al. (2007) CD133+ hepatic stellate cells are progenitor cells. *Biochem Biophys Res Commun* 52(2): 410-417.
3. Marsden ER, Hu Z, Fujio K, Nakatsukasa H, Thorgeirsson SS, Evarts RP (1992) Expression of acidic fibroblast growth factor in regenerating liver and during hepatic differentiation. *Lab Invest* 67(4): 427-433.
4. Schirmacher P, Geerts A, Pietrangelo A, Dienes HP, Rogler CE (1992) Hepatocyte growth factor/hepatopoietin A is expressed in fat-storing cells from rat liver but not myofibroblast-like cells derived from fat-storing cells. *Hepatology* 15: 5-11.
5. Maher J.J. (1993) Cell-specific expression of hepatocyte growth factor in liver. Upregulation in sinusoidal endothelial cells after carbon tetrachloride. *J Clin Invest* 91(5): 2244-2252.
6. Pinzani M, Knauss TC, Pierce GF, Hsieh P, Kenney W, Dubyak GR, Abboud HE (1991) Mitogenic signals for platelet-derived growth factor isoforms in liver fat-storing cells. *Am J Physiol* 260(3): C485-C491.

7. Higgins G.M., Anderson R.M. (1931) Experimental pathology of the liver:  
I. Restoration of the liver of the white rat following partial surgical removal.  
Arch Pathol 12: 186-202.
8. Gumerova AA, Valeeva IKh, Kiiasov AP. (1999) The "direct mitogen"  
lead nitrate causes an intensification in lipid peroxidation and Ito cell  
activation in the rat liver [Article in Russian] Ontogenez 30(4):289-295.