

My research interest branches into two related areas: (1) reprogramming somatic differentiated mammalian cells into stem-like cells through sphere formation in vitro, and re-directing them to specific differentiation for tissue regeneration in vivo; (2) analyzing the effects of epithelial-to-mesenchymal transition (EMT) transcription factor Zeb1 on transition of fibroblasts to sphere-derived stem-like cells (SDSC).

Tissue damages or functional disorders due to trauma, aging, and inheritable diseases like retinitis pigmentosa (RP), and age-related macular degeneration (AMD), are difficult to treat. More and more hope now relies on advances in regenerative medicine in which stem cell application is part of the solution. Recently, we have developed a novel protocol for reprogramming fibroblasts to immortal multipotential adult stem-like cells. This reprogramming pathway involves sequential mesenchymal-to-epithelial transition (MET), hypoxic induction of Aid and in turn Oct4 and Dnmt1-dependent silencing of cdk inhibitors and Arf to cause immortalization. We are hoping that application of SDSC that are not tumorigenic in vivo though immortal in vitro will facilitate patient-specific cell therapies in the clinic.

Publications

Liu Y, Mukhopadhyay P, Pisano MM, Lu X, Huang L, Lu Q, Dean DC. Repression of Zeb1 and Hypoxia Cause Sequential MET and Induction of Aid, Oct4, and Dnmt1, Leading to Immortalization and Multipotential Reprogramming of Fibroblasts in Spheres. Stem Cells. 2013 Apr 4. doi: 10.1002/stem.1382. [Epub ahead of print]

Liu Y, Sanchez-Tillo E, Lu X, Huang L, Clem B, Telang S, Jenson AB, Cuatrecasas M, Chesney J, Postigo A, Dean DC. Sequential Inductions of the ZEB1 Transcription Factor Caused by Mutation of Rb and then Ras are required for Tumor Initiation and Progression. J Biol Chem. 2013 Feb 26. [Epub ahead of print]

Reynolds M, Lane AN, Robertson B, Kemp S, **Liu Y**, Hill BG, Dean DC, Clem BF. Control of Glutamine Metabolism By the Tumor Suppressor Rb. Oncogene. doi: 0.1038/onc.2012.635. [Epub ahead of print]

Sánchez-Tilló E, **Liu Y**, de Barrios O, Siles L, Fanlo L, Cuatrecasas M, Darling DS, Dean DC, Castells A, Postigo A. EMT-activating transcription factors in cancer: beyond EMT and tumor invasiveness. Cell Mol Life Sci. 2012. 69(20):3429-56

Zhou L, Wang W, **Liu Y**, Fernandez de Castro J, Ezashi T, Telugu BP, Roberts RM, Kaplan HJ, Dean DC. Differentiation of Swine iPSC into Rod Photoreceptors and Their Integration into the Retina. Stem Cells. 29(6):972-80, 2011.



