

Hippo signaling in and around liver cancer

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The Hippo pathway effectors Yap and Taz are well-known drivers of tumor growth. For example, deletion of Yap/Taz in liver cancer in mice causes complete tumor regression. Unexpectedly, however, we found that Yap/Taz also exert a tumor suppressive function, where their activation in normal cells surrounding liver tumors suppresses tumor growth. Normal hepatocytes around liver tumors activated Yap/Taz and deletion of Yap/Taz in peritumoral hepatocytes accelerated tumor growth. Conversely, experimental hyperactivation of Yap in peritumoral hepatocytes triggered regression of primary liver tumors and melanoma-derived liver metastases. These data indicate that a major function of Yap/Taz in and around liver tumors is to promote cellular fitness and cell competition.

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