Akt/PKB-Mediated Phosphorylation of Twist1 Promotes Tumor Metastasis via Mediating Cross-Talk between PI3K/Akt and TGF-β Signaling Axes .................. 248

Précis: Phosphorylation of TWIST1 by AKT promotes EMT and metastasis via TGF-β2 transcriptional regulation and PI3K/AKT feedback activation.

nab-Paclitaxel Potentiates Gemcitabine Activity by Reducing Cytidine Deaminase Levels in a Mouse Model of Pancreatic Cancer . . . 260
K.K. Frese, A. Neesse, N. Cook, T.E. Bapiro, M.P. Lolkema, D.J. Jodrell, and D.A. Tuveson

Précis: Combined nab-paclitaxel and gemcitabine therapy leads to synergistic antitumor effects due to decreased gemcitabine metabolism.

ON THE COVER Frese and colleagues utilized a genetically engineered mouse model of pancreatic ductal adenocarcinoma (PDA) to better understand the mechanistic basis for the clinical observation that nab-paclitaxel, a water-soluble, albumin-bound form of paclitaxel, elicits synergistic antitumor activity when combined with gemcitabine, a nucleoside analogue that is the current standard of care for PDA. Combination treatment with nab-paclitaxel increases intratumoral gemcitabine levels by creating an oxidative environment within the tumor that promotes degradation of cytidine deaminase, the primary gemcitabine metabolizing enzyme. For details, please see the article by Frese and colleagues on page 260.

Suppression of Tumor Invasion and Metastasis by Concurrent Inhibition of c-Met and VEGF Signaling in Pancreatic Neuroendocrine Tumors ............... 270

Précis: Combined inhibition of VEGF and c-MET reduces the tumor invasiveness and metastasis observed after inhibition of VEGF alone and decreases tumor growth and angiogenesis.

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