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Remodeling of the Collagen Matrix in Aging Skin Promotes Melanoma Metastasis and Affects Immune Cell Motility ......................... 64
Précis: Loss of the cross-linking protein HAPLN1 in skin fibroblasts during aging results in changes to the extracellular matrix architecture that enhance melanoma cell invasion but impair immune cell infiltration.
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Age-Related Changes in HAPLN1 Increase Lymphatic Permeability and Affect Routes of Melanoma Metastasis .......................... 82
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Homophilic CD44 Interactions Mediate Tumor Cell Aggregation and Polyclonal Metastasis in Patient-Derived Breast Cancer Models .................................................. 96
Précis: Intercellular homophilic CD44 interaction-mediated tumor cell aggregation recruits PAK2 and activates the PAK2 signaling pathway to drive tumorigenesis and polyclonal metastasis.
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Spatiotemporal Loss of NF1 in Schwann Cell Lineage Leads to Different Types of Cutaneous Neurofibroma Susceptible to Modification by the Hippo Pathway ........ 114
Précis: Inactivation of NF1 in a Hoxb7-expressing cell lineage faithfully models neurofibromagenesis and provides insights into the developmental origin of both plexiform and cutaneous neurofibroma.

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Précis: Loss of NF1 in Prss56-expressing boundary cap cells recapitulates both cutaneous and plexiform neurofibroma and provides insight into neurofibromagenesis.

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ON THE COVER
Kitajima and colleagues found that KRAS-mutant/LKB1-null (KL) lung cancers exhibit upregulation of genes associated with oxidative stress and downregulation of genes involved in interferon signaling, including TMEM173, which encodes the dsDNA sensor STING. A screen of epigenetic inhibitors revealed that inhibiting DNMT1 and EZH2 restored STING expression in STING-low KL cell lines, and the binding of DNMT1 to the TMEM173 promoter was increased in STING-null KL cell lines compared to STING-low KL cell lines. Ectopic expression of LKB1 combined with inhibition of DNMT1 restored STING expression and pathway activation, and a STING agonist restored PD-L1 expression in KL cell lines. Collectively, these findings demonstrate that LKB1 negatively regulates STING in KRAS-driven lung cancer and suggest potential therapeutic strategies for patients with KRAS-mutant/LKB1-null lung cancer. For details, please see the article by Kitajima and colleagues on page 34.