The Androgen-Regulated Protease TMPRSS2 Activates a Proteolytic Cascade Involving Components of the Tumor Microenvironment and Promotes Prostate Cancer Metastasis .......... 1310

Précis: The serine protease TMPRSS2 enhances androgen-driven prostate cancer metastasis by inducing HGF cleavage and activation of c-MET signaling, and may represent a potential therapeutic target.

See commentary, p. 1262

The Genomic Landscape of Pediatric Ewing Sarcoma .......... 1326

Précis: The serine protease TMPRSS2 enhances androgen-driven prostate cancer metastasis by inducing HGF cleavage and activation of c-MET signaling, and may represent a potential therapeutic target.

Using next-generation sequencing, Crompton, Stewart, and colleagues found that Ewing sarcoma tumors were relatively genetically stable, but exhibited recurrent loss of stromal antigen 2 (STAG2) expression, which was associated with metastatic progression. In addition, relapsed tumors displayed an increased mutation rate compared with tumors at diagnosis. Using whole-genome sequencing, Tirode, Surdez, and colleagues also detected few somatic alterations at diagnosis but frequently exhibited loss of STAG2 expression, which is correlated with metastatic progression.

Précis: Ewing sarcoma tumors exhibit a low mutation rate but frequently harbor somatic mutations in STAG2, which are mutually exclusive with CDKN2A loss and correlate with TP53 mutations and poor prognosis.

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