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The Landscape of Human Cancer Proteins Targeted by SARS-CoV-2 916

B. Tutuncuoglu, M. Cakir, J. Batra, M. Bouhaddou, M. Eckhardt, D.E. Gordon, and N.J. Krogan

PRÉCIS: Patients with cancer in a New York hospital system were much more vulnerable to COVID-19 death than the general population, with a case fatality rate that varied by cancer type and was 28% overall.

IN THE SPOTLIGHT

Fusing the Genetic Landscape of Infantile High-Grade Gliomas 904
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Utility of Human-Derived Models for Glioblastoma 907
X. Luo and W.A. Weiss

It Takes a Village to Overcome KRAS Dependence in Pancreatic Cancer 910
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The Era of COVID-19 and the Rise of Science Collectivism in Cancer Research 913
T. Janowitcz and D.A. Tuveson

FOR MORE NEWS AND RESEARCH WATCH, VISIT CANCER DISCOVERY ONLINE AT http://cancerdiscovery.aacrjournals.org/CDNews.
A subset of infant high-grade gliomas with shared molecular and clinical characteristics were associated with better prognosis and often harbored targetable fusions involving genes encoding receptor tyrosine kinases.

See commentary, p. 904

**Tumor Microenvironment Is Critical for the Maintenance of Cellular States Found in Primary Glioblastomas**


Précis: Of four glioma stem cell–derived glioblastoma models, glioblastoma cerebral organoids most closely recapitulated the transcriptome and cell composition of primary tumors, a microenvironment-dependent effect.

See commentary, p. 907

**Gain-of-Function Genetic Alterations of G9a Drive Oncogenesis**


Précis: Amplification of or activating mutations in the histone methyltransferase–encoding gene EHMT2 reduced DKK1-mediated inhibition of the WNT pathway to promote melanoma development.

**EZH2-Deficient T-cell Acute Lymphoblastic Leukemia Is Sensitized to CHK1 Inhibition through Enhanced Replication Stress**


Précis: In T-cell acute lymphoblastic leukemia (T-ALL), loss-of-function mutations affecting EZH2 confer poor prognosis, but elevated replication stress may render EZH2-mutant T-ALL sensitive to CHK1 inhibition.

**Selective Alanine Transporter Utilization Creates a Targetable Metabolic Niche in Pancreatic Cancer**


Précis: Pancreatic ductal adenocarcinoma (PDAC) cells used the neutral amino acid transporter SLC38A2 to import necessary alanine, and lack of SLC38A2 caused a metabolic crisis in PDAC cells and tumor regression in vivo.

**Somatic Tissue Engineering in Mouse Models Reveals an Actionable Role for WNT Pathway Alterations in Prostate Cancer Metastasis**


Précis: A rapid, targeted method to generate genetically engineered mouse models of prostate cancer was developed and used to show that tankyrase inhibition may be useful in WNT pathway–activated disease.

**Tumor Microenvironment Remodeling Enables Bypass of Oncogenic KRAS Dependency in Pancreatic Cancer**


Précis: Pancreatic ductal adenocarcinoma escaped dependence on oncogenic KRAS via an HDAC5-mediated mechanism that resulted in macrophage recruitment to the tumor microenvironment via a druggable pathway.

See commentary, p. 910

**Correction**

Correction: Targeting HER2 with Trastuzumab Deruxtecan: A Dose-Expansion, Phase I Study in Multiple Advanced Solid Tumors ............... 1078
Because no model is perfect, Pine, Cirigliano, and colleagues sought to characterize four commonly used glioblastoma models: two-dimensional glioma sphere cultures, three-dimensional tumor organoids, glioblastoma cerebral organoids (GLICO), and patient-derived xenografts. GLICOs stood out as most closely resembling primary glioblastomas in several important ways, with closely overlapping transcriptomes and similarities in cell-type composition. GLICOs’ ability to recapitulate many aspects of glioblastoma biology depended on the microenvironment: When cultured in two-dimensional conditions, GLICO-derived cells lost many similarities with primary glioblastomas. This work showcases the strengths of GLICOs and provides detailed characterizations of the three other models, providing researchers with data to make informed decisions about which model best suits their purposes. For more information, see the article by Pine, Cirigliano, and colleagues on page 964.