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The Landscape of Human Cancer Proteins Targeted by SARS-CoV-2 ...............916
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RESEARCH BRIEF
Case Fatality Rate of Cancer Patients with COVID-19 in a New York Hospital System . . .935

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.

RESEARCH ARTICLES

Infant High-Grade Gliomas Comprise Multiple Subgroups Characterized by Novel Targetable Gene Fusions and Favorable Outcomes ...............942
Tumor Microenvironment Is Critical for the Maintenance of Cellular States Found in Primary Glioblastomas

Precis: 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

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

See commentary, p. 910

Correction

Correction: Targeting HER2 with Trastuzumab Deruxtecan: A Dose-Expansion, Phase I Study in Multiple Advanced Solid Tumors

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

Precis: 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

Precis: 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

Precis: 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
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.