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1824 Second-Line Therapies in Hepatocellular Carcinoma: Emergence of Resistance to Sorafenib Augusto Villanueva and Josep M. Llovet
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1830 New Strategies for Advanced Neuroendocrine Tumors in the Era of Targeted Therapy Mei Dong, Alexandria T. Phan, and James C. Yao

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HUMAN CANCER BIOLOGY

Immune Thrombocytopenia in Patients with Chronic Lymphocytic Leukemia Is Associated with Stereotyped B-cell Receptors Carlo Visco, Francesco Maura, Giacomo Tuana, Luca Agnelli, Marta Lionetti, Sonia Fabris, Elisabetta Novella, Ilaria Giaretta, Gianluigi Reda, Wilma Barcellini, Luca Baldini, Antonino Neri, Francesco Rodeghiero, and Agostino Cortelezz

MAPKAP Kinase 2 Overexpression Influences Prognosis in Gastrointestinal Stromal Tumors and Associates with Copy Number Variations on Chromosome 1 and Expression of p58 MAP Kinase and ET1
Peter Birner, Andrea Beer, Ursula Vinatzer, Susanne Starzy, Romana Höflberger, Nadine Nirtl, Fritz Wrb, Berthold Streubel, and Sebastian F. Schoppmann

Protein Kinase CK2 Protects Multiple Myeloma Cells from ER Stress–Induced Apoptosis and from the Cytotoxic Effect of HSP90 Inhibition through Regulation of the Unfolded Protein Response Sabrina Manni, Alessandra Brancalion, Laura Quotti Tubi, Anna Colpo, Laura Pavan, Anna Cabrelle, Elisa Ave, Fortunato Zaffino, Giovanni Di Maira, Maria Ruzzene, Fausto Adami, Renato Zambello, Maria Rita Pitari, Pierfrancesco Tassone, Lorenzo A. Pinna, Carmela Gurrieri, Gianpietro Semenzato, and Francesco Piazza

Glioblastoma Stem–like Cell Lines with Either Maintenance or Loss of High-Level EGFR Amplification, Generated via Modulation of Ligand Concentration Alexander Schulte, Hauke S. Günther, Tobias Martens, Svenja Zapf, Sabine Riethdorf, Clemens Wülfing, Malgorzata Stoupiec, Manfred Westphal, and Katrin Lamszus
1914 L1 Cell Adhesion Molecule Promotes Tumorigenicity and Metastatic Potential in Non–Small Cell Lung Cancer
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1925 Immune Suppression in Premalignant Respiratory Papillomas: Enriched Functional CD4⁺Foxp3⁺ Regulatory T Cells and PD-1/PD-L1/L2 Expression
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1936 Integrative Genomics Identified RFC3 As an Amplified Candidate Oncogene in Esophageal Adenocarcinoma

1947 Frequency of Driver Mutations in Lung Adenocarcinoma from Female Never-Smokers Varies with Histologic Subtypes and Age at Diagnosis
Yang Zhang, Yihua Sun, Yunjian Pan, Chenguang Li, Lei Shen, Yuan Li, Xiaoyang Luo, Ting Ye, Rui Wang, Haichuan Hu, Hang Li, Lei Wang, William Pao, and Haiquan Chen

1992 Serum Autoantibody Signature of Ductal Carcinoma In Situ Progression to Invasive Breast Cancer
Alain Mangé, Jérôme Lacombe, Caroline Bascoul-Mollevi, Marta Jarlier, Pierre-Jean Lamy, Philippe Rouanet, Thierry Maudelonde, and Jérôme Solassol

2001 Copy Number Gain of 1q25 Predicts Poor Progression-Free Survival for Pediatric Intracranial Ependymomas and Enables Patient Risk Stratification: A Prospective European Clinical Trial Cohort Analysis on Behalf of the Children’s Cancer Leukaemia Group (CCLG), Société Française d’Oncologie Pédiatrique (SFOP), and International Society for Pediatric Oncology (SIOP)

2012 A Three-Gene Expression Signature Model for Risk Stratification of Patients with Neuroblastoma
Idoia Garcia, Gemma Mayol, José Rios, Gema Domenech, Nai-Kong V. Cheung, André Oberthuer, Matthias Fischer, John M. Maris, Garrett M. Brodeur, Barbara Hero, Eva Rodríguez, Mariona Suñol, Patricia Galvan, Carmen de Torres, Jaume Mora, and Cinzia Lavarino

2024 ¹⁸F-FDG-PET/CT Imaging as an Early Survival Predictor in Patients with Primary High-Grade Soft Tissue Sarcomas Undergoing Neoadjuvant Therapy
A Panel of Four miRNAs Accurately Differentiates Malignant from Benign Indeterminate Thyroid Lesions on Fine Needle Aspiration
Xavier M. Keutgen, Filippo Filicori, Michael J. Crowley, Yongchun Wang, Theresa Scognamiglio, Rana Hoda, Daniel Buitrago, David Cooper, Martha A. Zeiger, Rasa Zarnegar, Olivier Elemento, and Thomas J. Fahey III

CTLA-4 Blockade with Ipilimumab: Long-term Follow-up of 177 Patients with Metastatic Melanoma
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Phase I Study of Rigosertib, an Inhibitor of the Phosphatidylinositol 3-Kinase and Polo-like Kinase 1 Pathways, Combined with Gemcitabine in Patients with Solid Tumors and Pancreatic Cancer
Wen Wee Ma, Wells A. Messersmith, Grace K. Dy, Colin D. Weekes, Amy Whitworth, Chen Ren, Manoj Maniar, Francois Wilhelm, S. Gail Eckhardt, Alex A. Adjei, and Antonio Jimeno

Phase II Efficacy and Pharmacogenomic Study of Selumetinib (AZD6244; ARRY-142886) in Iodine-131 Refractory Papillary Thyroid Carcinoma with or without Follicular Elements
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Implications of Plasma Protein Binding for Pharmacokinetics and Pharmacodynamics of the γ-Secretase Inhibitor RO4929097
Jianmei Wu, Patricia M. LoRusso, Larry H. Matherly, and Jing Li

Phase I/II Trial of Carboplatin and Paclitaxel Chemotherapy in Combination with Intravenous Oncolytic Reovirus in Patients with Advanced Malignancies
Eleni M. Karapanagiotou, Victoria Roulstone, Katie Twigger, Merced Ball, MaryAnne Tanay, Chris Nutting, Kate Newbold, Martin E. Gore, James Larkin, Konstantinos N. Syrigos, Matt Coffey, Brad Thompson, Karl Mettinger, Richard G. Vile, Hardev S. Pandha, Geoff D. Hall, Alan A. Melcher, John Chester, and Kevin J. Harrington

Phase II, Open-Label Study of Brivanib as Second-Line Therapy in Patients with Advanced Hepatocellular Carcinoma
Richard S. Finn, Yoon-Koo Kang, Mary Mulcahy, Blase N. Politi, Ho Yeong Lim, Ian Walters, Christine Baudelet, Demetrios Manekas, and Joong-Won Park
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Sorafenib Is an Inhibitor of UGT1A1 but Is Metabolized by UGT1A9: Implications of Genetic Variants on Pharmacokinetics and Hyperbilirubinemia
Cody J. Peer, Tristan M. Sissung, AeRang Kim, Lokesh Jain, Sukyung Woo, Erin R. Gardner, C. Tyler Kirkland, Sarah M. Troutman, Bevin C. English, Emily D. Richardson, Joel Feder Spielberg, David Venzon, William D. Figg

Tumor Hypoxia Predicts Biochemical Failure following Radiotherapy for Clinically Localized Prostate Cancer
Michael Milosevic, Padraig Warde, Cynthia Ménard, Peter Chung, Ants Toi, Adrian Ishkanian, Michael McLean, Melanie Pinililé, Jemma Sykes, Mary Gospodarowicz, Charles Catton, Richard P. Hill, and Robert Bristow

Correction: Molecular Imaging of TGFβ-Induced Smad2/3 Phosphorylation Reveals a Role for Receptor Tyrosine Kinases in Modulating TGFβ Signaling

PREDICTIVE BIOMARKERS AND PERSONALIZED MEDICINE
ABOUT THE COVER

High-level EGFR gene amplification can be retained in glioblastoma stem-like cell lines established and propagated without recombinant EGF. In contrast, high-level amplification is lost in parallel cell lines from the same tumors established with EGF supplementation. Cell lines with high-level EGFR amplification produce highly aggressive xenograft tumors in the brains of nude mice, retaining the EGFR amplification as shown in the cover figure, whereas counterpart cell lines, lacking high-level amplification, are either nontumorigenic or grow significantly more slowly in vivo. For details, see the article by Schulte and colleagues on page 1901 of this issue.