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November 15, 2010 • Volume 70 • Number 22

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Meghan J. Seltzer, Bryson D. Bennett, Avadhut D. Joshi, Ping Gao, Ajit G. Thomas, Dana V. Ferraris, Takashi Tsukamoto, Camilo J. Rojas, Barbara S. Slusher, Joshua D. Rabinowitz, Chi V. Dang, and Gregory J. Riggins

**Précis:** IDH1 mutations create a higher dependence of glioma cells on  $\alpha$ -ketoglutarate, relative to normal cells, that could be therapeutically exploited by targeting glutamine metabolism.

8988

Heat Shock Protein 60 Regulation of the Mitochondrial Permeability Transition Pore in Tumor Cells

Jagadish C. Ghosh, Markus D. Siegelin, Takehiko Dohi, and Dario C. Altieri

**Précis:** Findings implicate a chaperone network in controlling mitochondrial permeability transition and apoptosis in cancer cells.

#### **CLINICAL STUDIES**

8994

The Ability of Biomarkers to Predict Systemic Progression in Men with High-Risk Prostate Cancer Treated Surgically Is Dependent on ERG Status

R. Jeffrey Karnes, John C. Cheville, Cristiane M. Ida, Thomas J. Sebo, Asha A. Nair, Hui Tang, Jan-Marie Munz, Farhad Kosari, and George Vasmatzis

**Précis:** The study presents TOP2A as a superior prognostic marker for prostate cancer and shows that its predictive ability is dependent on cancer subtype.

9003

Repeat Dose Study of the Cancer Chemopreventive Agent Resveratrol in Healthy Volunteers: Safety, Pharmacokinetics, and Effect on the Insulin-like Growth Factor Axis

Victoria A. Brown, Ketan R. Patel, Maria Viskaduraki, James A. Crowell, Marjorie Perloff, Tristan D. Booth, Grygoriy Vasilinin, Ananda Sen, Anna Maria Schinas, Gianfranca Piccirilli, Karen Brown, William P. Steward, Andreas J. Gescher, and Dean E. Brenner

**Précis:** This study defines suitable doses, accompanying blood levels, and potential efficacy biomarkers which might help to optimize future cancer chemoprevention intervention studies of the red wine constituent resveratrol.

### INTEGRATED SYSTEMS AND TECHNOLOGIES

9012

Multimodal Mass Spectrometric Imaging of Small Molecules Reveals Distinct Spatio-Molecular Signatures in Differentially Metastatic Breast Tumor Models

Erika R. Amstalden van Hove, Tiffany R. Blackwell, Ivo Klinkert, Gert B. Eijkel, Ron M.A. Heeren, and Kristine Glunde

**Précis:** Mass spectrometry imaging techniques can identify different tissue microenvironments in breast tumors, possibly improving their clinicopathological assessment in the future.

9022

PET Imaging of Tumor Neovascularization in a Transgenic Mouse Model with a Novel <sup>64</sup>Cu-DOTA-Knottin Peptide

Carsten H. Nielsen, Richard H. Kimura, Nadia Withofs, Phuoc T. Tran, Zheng Miao, Jennifer R. Cochran, Zhen Cheng, Dean Felsher, Andreas Kjær, Juergen K. Willmann, and Sanjiy S. Gambhir

**Précis:** Methods that improve PET imaging in cancer offer opportunities for improved detection and management.

### MICROENVIRONMENT AND IMMUNOLOGY

9031

Intranodal Vaccination with Naked Antigen-Encoding RNA Elicits Potent Prophylactic and Therapeutic Antitumoral Immunity

Sebastian Kreiter, Abderraouf Selmi, Mustafa Diken, Michael Koslowski, Cedrik M. Britten, Christoph Huber, Özlem Türeci, and Ugur Sahin

**Précis:** Findings define methods to achieve strong antigen-specific immunity and cancer cures using antigen-encoding RNA in preclinical animal models.

9041

A Therapeutic OX40 Agonist Dynamically Alters Dendritic, Endothelial, and T Cell Subsets within the Established Tumor Microenvironment

Angela D. Pardee, Dustin McCurry, Sean Alber, Peisheng Hu, Alan L. Epstein, and Walter J. Storkus

**Précis:** Correcting tumoral immune escape exerts therapeutic effects against established tumors by improving the delivery of antitumor T cells into the tumor microenvironment.

9053

Multiple Injections of Electroporated Autologous T Cells Expressing a Chimeric Antigen Receptor Mediate Regression of Human Disseminated Tumor

Yangbing Zhao, Edmund Moon, Carmine Carpenito, Chrystal M. Paulos, Xiaojun Liu, Andrea L. Brennan, Anne Chew, Richard G. Carroll, John Scholler, Bruce L. Levine, Steven M. Albelda, and Carl H. June

**Précis:** Findings offer preclinical proof-of-concept for a novel adoptive T-cell transfer approach for cancer treatment that can potentiate antitumor effects and complement approaches using retroviral and lentiviral engineered T-cell therapies.

9062

Improved Clinical Outcome in Indolent B-Cell Lymphoma Patients Vaccinated with Autologous Tumor Cells Experiencing Immunogenic Death

Roberta Zappasodi, Serenella M. Pupa, Gaia C. Ghedini, Italia Bongarzone, Michele Magni, Antonello D. Cabras, Mario P. Colombo, Carmelo Carlo-Stella, Alessandro M. Gianni, and Massimo Di Nicola

**Précis:** Clinical findings confirm expectations of a stringent correlation between the therapeutic efficacy of a dendritic cell vaccine and the extent of immunogenic death induced in the tumor cells used to load the dendritic cells before vaccination.

9073

High-Throughput Characterization of 10 New Minor Histocompatibility Antigens by Whole Genome Association Scanning

Cornelis A.M. Van Bergen, Caroline E. Rutten, Edith D. Van Der Meijden, Simone A.P. Van Luxemburg-Heijs, Ellie G.A. Lurvink, Jeanine J. Houwing-Duistermaat, Michel G.D. Kester, Arend Mulder, Roel Willemze, J.H. Frederik Falkenburg, and Marieke Griffioen

**Précis:** In this study, efficient T-cell selection was combined with SNP genotyping to discover a significant number of new minor histocompatibility antigens that can be targeted by cellular immunotherapy of cancer.

### MOLECULAR AND CELLULAR PATHOBIOLOGY

9084

#### Critical Roles of DMP1 in Human Epidermal Growth Factor Receptor 2/neu-Arf-p53 Signaling and Breast Cancer Development

Pankaj Taneja, Dejan Maglic, Fumitake Kai, Takayuki Sugiyama, Robert D. Kendig, Donna P. Frazier, Mark C. Willingham, and Kazushi Inoue

**Précis:** Mechanistic findings define a transcriptional cascade that plays a critical role in quenching HER2/neu oncogenic signals known to promote breast cancer.

9095

#### Global Downstream Pathway Analysis Reveals a Dependence of Oncogenic NF-E2-Related Factor 2 Mutation on the mTOR Growth Signaling Pathway

Tatsuhiro Shibata, Shigeru Saito, Akiko Kokubu, Takafumi Suzuki, Masayuki Yamamoto, and Setsuo Hirohashi

**Précis:** Findings offer a rationale to target a master transcriptional regulator of antioxidant genes and phase II detoxifying enzymes as an anticancer strategy, also suggesting it as a novel theranostic biomarker.

9106

#### Insulin-Like Growth Factor I Suppresses Bone Morphogenetic Protein Signaling in Prostate Cancer Cells by Activating mTOR Signaling

Reema S. Wahdan-Alaswad, Kyung Song, Tracy L. Krebs, Dorjee T.N. Shola, Jose A. Gomez, Shigemi Matsuyama, and David Danielpour

Précis: Findings elucidate how a key growth factor in prostate cancer acts at several levels to shut down a central pathway of epithelial cell function in the prostate, with important therapeutic implications for prostate cancer management.

9118

#### The Mitotic Kinase Aurora-A Induces Mammary Cell Migration and Breast Cancer Metastasis by Activating the Cofilin-F-actin Pathway

Li-hui Wang, Jin Xiang, Min Yan, Yan Zhang, Yan Zhao, Cai-feng Yue, Jie Xu, Fei-meng Zheng, Jin-na Chen, Zhuang Kang, Tong-sheng Chen, Da Xing, and Quentin Liu

**Précis:** An important mitotic kinase that is targeted by small molecule inhibitors currently in clinical testing is found to have an important role in supporting metastasis.

9129

#### Metabolic Profiling Detects Field Effects in Nondysplastic Tissue from Esophageal Cancer Patients

Danny Yakoub, Hector C. Keun, Robert Goldin, and George B. Hanna

**Précis:** Esophageal cancer can be detected by metabolic profiling of adjacent histologically normal mucosa.

9137

#### Dysregulation of the Mitogen Granulin in Human Cancer through the miR-15/107 microRNA Gene Group

Wang-Xia Wang, Natasha Kyprianou, Xiaowei Wang, and Peter T. Nelson

**Précis:** Findings elucidate microRNA-mediated control of a common mitogen in a variety of human cancers, the regulation of which had been previously obscure, with implications for microRNA-based therapeutic targeting.

9143

#### Mammalian Target of Rapamycin–Dependent Acinar Cell Neoplasia after Inactivation of *Apc* and *Pten* in the Mouse Salivary Gland: Implications for Human Acinic Cell Carcinoma

Cassandra R. Diegel, Kathleen R. Cho, Adel K. El-Naggar, Bart O. Williams, and Charlotta Lindvall

**Précis:** Findings report a novel preclinical model and a rationale to evaluate rapamycin therapy for salivary gland acinic cell carcinoma, a low grade neoplasm but one that tends to recur.

9153

#### Expression Profiling in Progressive Stages of Fumarate-Hydratase Deficiency: The Contribution of Metabolic Changes to Tumorigenesis

Houman Ashrafian, Linda O'Flaherty,
Julie Adam, Violetta Steeples, Yuen-Li Chung,
Phil East, Sakari Vanharanta, Heli Lehtonen,
Emma Nye, Emine Hatipoglu, Melroy Miranda,
Kimberley Howarth, Deepa Shukla, Helen Troy,
John Griffiths, Bradley Spencer-Dene,
Mohammed Yusuf, Emanuela Volpi,
Patrick H. Maxwell, Gordon Stamp,
Richard Poulsom, Christopher W. Pugh,
Barbara Costa, Chiara Bardella,
Maria Flavia Di Renzo, Michael I. Kotlikoff,
Virpi Launonen, Lauri Aaltonen,
Mona El-Bahrawy, Ian Tomlinson,
and Patrick J. Pollard

**Précis:** Fumarate hydratase deficiency compels cells to prematurely adopt a metabolic phenotype that predisposes them to cell proliferation and growth and ultimately to cancer.

#### Acquired TNFRSF14 Mutations in Follicular Lymphoma Are Associated with Worse Prognosis

K-John J. Cheung, Nathalie A. Johnson, Joslynn G. Affleck, Tesa Severson, Christian Steidl, Susana Ben-Neriah, Jacqueline Schein, Ryan D. Morin, Richard Moore, Sohrab P. Shah, Hong Qian, Jessica E. Paul, Adele Telenius, Thomas Relander, Wan Lam, Kerry Savage, Joseph M. Connors, Carolyn Brown, Marco A. Marra, Randy D. Gascoyne, and Douglas E. Horsman

**Précis:** Findings identify a candidate genetic marker of prognosis in follicular lymphoma, which may provide an explanation as to why its treatment with rituximab provides less benefit in some patients.

9175

## Epigenetically Deregulated microRNA-375 Is Involved in a Positive Feedback Loop with Estrogen Receptor $\alpha$ in Breast Cancer Cells

Pedro de Souza Rocha Simonini, Achim Breiling, Nibedita Gupta, Mahdi Malekpour, Mahmoud Youns, Ramesh Omranipour, Fatemeh Malekpour, Stefano Volinia, Carlo M. Croce, Hossein Najmabadi, Sven Diederichs, Özgür Sahin, Doris Mayer, Frank Lyko, Jörg D. Hoheisel, and Yasser Riazalhosseini

**Précis:** Findings offer significant new insights into how the estrogen receptor is deregulated in breast cancer.

9185

#### Alternative Splicing of Caspase 9 Is Modulated by the Phosphoinositide 3-Kinase/Akt Pathway via Phosphorylation of SRp30a

Jacqueline C. Shultz, Rachel W. Goehe, D. Shanaka Wijesinghe, Charuta Murudkar, Amy J. Hawkins, Jerry W. Shay, John D. Minna, and Charles E. Chalfant

Précis: Findings define a functionally important alternative splicing mechanism in lung adenocarcinoma that offers a novel approach to therapeutic development.

9197

#### Upregulation of DLX5 Promotes Ovarian Cancer Cell Proliferation by Enhancing IRS-2-AKT Signaling

Yinfei Tan, Mitchell Cheung, Jianming Pei, Craig W. Menges, Andrew K. Godwin, and Joseph R. Testa

**Précis:** Findings define a homeobox protein that positively modifies a variety of oncogenic signals in ovarian cancer, with potential prognostic applications.

9207

#### Human RecQL4 Helicase Plays Critical Roles in Prostate Carcinogenesis

Yanrong Su, Jarah A. Meador, Gloria M. Calaf, Luca Proietti De-Santis, Yongliang Zhao, Vilhelm A. Bohr, and Adayabalam S. Balajee

**Précis:** Study implicates a DNA helicase regulated by Rb-E2F as an important driver of metastatic progression of prostate cancer.

#### PREVENTION AND EPIDEMIOLOGY

9218

#### Serum Vitamin D and Risk of Bladder Cancer

Alison M. Mondul, Stephanie J. Weinstein, Satu Männistö, Kirk Snyder, Ronald L. Horst, Jarmo Virtamo, and Demetrius Albanes

**Précis:** Accumulating evidence argues that serum levels of vitamin D exert a profound influence on cancer risk, as illustrated in this study of bladder cancer.

9224

#### Pesticide Use Modifies the Association Between Genetic Variants on Chromosome 8q24 and Prostate Cancer

Stella Koutros, Laura E. Beane Freeman, Sonja I. Berndt, Gabriella Andreotti, Jay H. Lubin, Dale P. Sandler, Jane A. Hoppin, Kai Yu, Qizhai Li, Laura A. Burdette, Jeffrey Yuenger, Meredith Yeager, and Michael C.R. Alavanja

**Précis:** Study offers an excellent illustration of how an environmental modifier can radically alter the impact of a genetic risk marker on a patient's susceptibilty to disease.

### THERAPEUTICS, TARGETS, AND CHEMICAL BIOLOGY

9234

#### Ki-67 as a Molecular Target for Therapy in an *In vitro* Three-Dimensional Model for Ovarian Cancer

Ramtin Rahmanzadeh, Prakash Rai, Jonathan P. Celli, Imran Rizvi, Bettina Baron-Lühr, Johannes Gerdes, and Tayyaba Hasan

**Précis:** A common immunohistopathological marker used for staging of clinical cancers may offer unexpected opportunities as a therapeutic target.

#### MGMT-Independent Temozolomide Resistance in Pediatric Glioblastoma Cells Associated with a PI3-Kinase–Mediated *HOX/*Stem Cell Gene Signature

Nathalie Gaspar, Lynley Marshall, Lara Perryman, Dorine A. Bax, Suzanne E. Little, Marta Viana-Pereira, Swee Y. Sharp, Gilles Vassal, Andrew D.J. Pearson, Rui M. Reis, Darren Hargrave, Paul Workman, and Chris Jones

**Précis:** Findings identify a reversible novel mechanism of resistance to the alkylating agent temozolomide in pediatric glioblastoma model systems and patient samples.

9253

#### Class III \( \beta\)-Tubulin Expression Predicts Prostate Tumor Aggressiveness and Patient Response to Docetaxel-Based Chemotherapy

Guillaume Ploussard, Stéphane Terry,
Pascale Maillé, Yves Allory, Nanor Sirab,
Laurence Kheuang, Pascale Soyeux,
Nathalie Nicolaiew, Estelle Coppolani,
Bernard Paule, Laurent Salomon,
Stéphane Culine, Ralph Buttyan,
Francis Vacherot, and Alexandre de la Taille

**Précis:** Findings suggest a novel theranostic molecule in advanced prostate cancer to predict responses to taxane therapy and improve treatment of taxane-resistant disease.

9265

#### Deubiquitinase Inhibition by Small-Molecule WP1130 Triggers Aggresome Formation and Tumor Cell Apoptosis

Vaibhav Kapuria, Luke F. Peterson, Dexing Fang, William G. Bornmann, Moshe Talpaz, and Nicholas J. Donato

**Précis:** An antitumor compound previously found to block signaling by Jak2 kinase is found to target deubiquitinases that are commonly overexpressed in cancer cells.

9277

#### Selective Killing of Tumor Neovasculature Paradoxically Improves Chemotherapy Delivery to Tumors

Freddy E. Escorcia, Erik Henke, Michael R. McDevitt, Carlos H. Villa, Peter Smith-Jones, Ronald G. Blasberg, Robert Benezra, and David A. Scheinberg

**Précis:** A novel approach to remodel the tumor vasculature has strongly positive pharmacokinetic consequences on cancer drug delivery in a preclinical model, resulting in better therapy and survival outcomes.

9287

#### Nox2 NADPH Oxidase Promotes Pathologic Cardiac Remodeling Associated with Doxorubicin Chemotherapy

Youyou Zhao, Declan McLaughlin, Emma Robinson, Adam P. Harvey, Michelle B. Hookham, Ajay M. Shah, Barbara J. McDermott, and David J. Grieve

**Précis:** A major ROS-producing enzyme in the heart is implicated in adverse cardiac remodeling caused by doxorubicin chemotherapy, implying a new therapeutic strategy to reduce doxorubicin cardiotoxicity.

9298

# WNT11 Expression Is Induced by Estrogen-Related Receptor $\alpha$ and $\beta$ -Catenin and Acts in an Autocrine Manner to Increase Cancer Cell Migration

Mary A. Dwyer, James D. Joseph, Hilary E. Wade, Matthew L. Eaton, Rebecca S. Kunder, Dmitri Kazmin, Ching-yi Chang, and Donald P. McDonnell

Précis: Results define an autocrine mechanism controlled by the nuclear receptor ERRα, associated with poor outcomes in several cancers, that influences cancer cell migratory capacity.

9309

#### Induction of DNA Damage-Inducible Gene GADD45β Contributes to Sorafenib-Induced Apoptosis in Hepatocellular Carcinoma Cells

Da-Liang Ou, Ying-Chun Shen, Sung-Liang Yu, Kuen-Feng Chen, Pei-Yen Yeh, Hsiang-Hsuan Fan, Wen-Chi Feng, Ching-Tzu Wang, Liang-In Lin, Chiun Hsu, and Ann-Lii Cheng

**Précis:** Mechanistic findings suggest a potential surrogate marker that could be useful to predict the efficacy of Sorafenib in liver cancer.

9319

#### Synergistic Enhancement of Carboplatin Efficacy with Photodynamic Therapy in a Three-Dimensional Model for Micrometastatic Ovarian Cancer

Imran Rizvi, Jonathan P. Celli, Conor L. Evans, Adnan O. Abu-Yousif, Alona Muzikansky, Brian W. Pogue, Dianne Finkelstein, and Tayyaba Hasan

Précis: A 3D model for ovarian micrometastases is introduced as a high-throughput platform to report treatment response, exemplified in this study to show how photodynamic therapy synergistically enhances carboplatin efficacy.

Voluntary Running Prevents Progressive Memory Decline and Increases Adult Hippocampal Neurogenesis and Growth Factor Expression after Whole-Brain Irradiation

Sarah J.E. Wong-Goodrich, Madeline L. Pfau, Catherine T. Flores, Jennifer A. Fraser, Christina L. Williams, and Lee W. Jones

**Précis:** Studies in mice suggest that voluntary exercise can alleviate progressive memory loss after cranial irradiation, strongly prompting studies in clinical settings.

9339

Telomerase-Dependent Oncolytic Adenovirus Sensitizes Human Cancer Cells to Ionizing Radiation via Inhibition of DNA Repair Machinery

Shinji Kuroda, Toshiya Fujiwara, Yasuhiro Shirakawa, Yasumoto Yamasaki, Shuya Yano, Futoshi Uno, Hiroshi Tazawa, Yuuri Hashimoto, Yuichi Watanabe, Kazuhiro Noma, Yasuo Urata, Shunsuke Kagawa, and Toshiyoshi Fujiwara

**Précis:** Results illustrate the potential of combining engineered oncolytic virotherapy and ionizing radiation as a strategy to improve the management of human cancer.

#### **TUMOR AND STEM CELL BIOLOGY**

9349

Brick1 Is an Essential Regulator of Actin Cytoskeleton Required for Embryonic Development and Cell Transformation

Beatriz Escobar, Guillermo de Cárcer, Gonzalo Fernández-Miranda, Alberto Cascón, José J. Bravo-Cordero, María C. Montoya, Mercedes Robledo, Marta Cañamero, and Marcos Malumbres

**Précis:** Genetic ablation of Brick1, a component of the Wave/Scar actin regulatory complex, appears to exert a protective effect against malignant development in renal cancer and other cancers where loss occurs.

9360

mTOR Complex Component Rictor Interacts with PKCζ and Regulates Cancer Cell Metastasis

Fei Zhang, Xiaofang Zhang, Menghui Li, Peng Chen, Bin Zhang, Hua Guo, Wenfeng Cao, Xiying Wei, Xuchen Cao, Xishan Hao, and Ning Zhang

**Précis:** Findings illuminate understanding of how EGF activates the mTORC2 growth regulatory complex to drive breast cancer metastasis, with implications for clinical studies of mTOR inhibitors.

9371

Surface-Immobilized Aptamers for Cancer Cell Isolation and Microscopic Cytology

Yuan Wan, Young-tae Kim, Na Li, Steve K. Cho, Robert Bachoo, Andrew D. Ellington, and Samir M. Iqbal

**Précis:** More sensitive and specific tumor detection technologies are needed to rapidly assess surgical resection margins and to monitor residual tumor cells that may persist in blood after disease treatment.

9381

Intrinsic Cooperation between p16 $^{\rm INK4a}$  and p21 $^{\rm Waf1/Cip1}$  in the Onset of Cellular Senescence and Tumor Suppression *In vivo* 

Shinji Takeuchi, Akiko Takahashi, Noriko Motoi, Shin Yoshimoto, Tomoko Tajima, Kimi Yamakoshi, Atsushi Hirao, Shigeru Yanagi, Kiyoko Fukami, Yuichi Ishikawa, Saburo Sone, Eiji Hara, and Naoko Ohtani

**Précis:** Expectations that mice lacking both the p16INK4 and p2IWAF1 tumor suppressor genes would be highly susceptible to carcinogenesis is experimentally confirmed and mechanistically dissected in this study.

9391

Tyrosine Phosphorylation Profiling Reveals the Signaling Network Characteristics of Basal Breast Cancer Cells

Falko Hochgräfe, Luxi Zhang, Sandra A. O'Toole, Brigid C. Browne, Mark Pinese, Ana Porta Cubas, Gillian M. Lehrbach, David R. Croucher, Danny Rickwood, Alice Boulghourjian, Robert Shearer, Radhika Nair, Alexander Swarbrick, Dana Faratian, Peter Mullen, David J. Harrison, Andrew V. Biankin, Robert L. Sutherland, Mark J. Raftery, and Roger J. Daly

**Précis:** Phosphoproteomic profiling reveals that basal breast cancer cells are characterized by an extensive Src family kinase signalling network, highlighting candidate theranostic targets for this aggressive subtype of breast cancers.

ANCCA/ATAD2 Overexpression Identifies Breast Cancer Patients with Poor Prognosis, Acting to Drive Proliferation and Survival of Triple-Negative Cells through Control of B-Myb and EZH2

Ekaterina V. Kalashnikova, Alexey S. Revenko, Abigael T. Gemo, Nicolas P. Andrews, Clifford G. Tepper, June X. Zou, Robert D. Cardiff, Alexander D. Borowsky, and Hong-Wu Chen

**Précis:** Findings suggest that the prognostic significance of a chromatin coregulator for hormone receptors in breast cancer is based on its role in integrating multiple oncogenic programs, with particular importance to aggressive triple-negative tumors.

9413

Quantitative *In vivo* Imaging of the Effects of Inhibiting Integrin Signaling via Src and FAK on Cancer Cell Movement: Effects on E-cadherin Dynamics

Marta Canel, Alan Serrels, Derek Miller, Paul Timpson, Bryan Serrels, Margaret C. Frame, and Valerie G. Brunton

**Précis:** Results identify a novel role for FAK and Src in regulating tumor cell movement in vivo using fluorescence based microscopy.

9423

Hepatocyte Nuclear Factor-4α Promotes Gut Neoplasia in Mice and Protects against the Production of Reactive Oxygen Species

Mathieu Darsigny, Jean-Philippe Babeu, Ernest G. Seidman, Fernand-Pierre Gendron, Emile Levy, Julie Carrier, Nathalie Perreault, and François Boudreau

Précis: Findings point to the nuclear receptor HNF4\(\mathbb{A}\) as a potential therapeutic target to modify epithelial cell resistance to ROS production during intestinal tumorigenesis.

9434

Clld7, A Candidate Tumor Suppressor on Chromosome 13q14, Regulates Pathways of DNA Damage/Repair and Apoptosis

Xiaobo Zhou and Karl Münger

**Précis:** Findings support the definition of a new tumor suppressor located at chromosome 13q14 and suggest its possible utility in predicting cancer chemosensitivity.

9444

Tumor-Initiating Function of Nucleostemin-Enriched Mammary Tumor Cells

Tao Lin, Lingjun Meng, Yi Li, and Robert Y.L. Tsai

**Précis:** Studies of a stem cell marker in breast cancer cells suggest its potential as a therapeutic target.

9453

Aberrant Activation of Fatty Acid Synthesis Suppresses Primary Cilium Formation and Distorts Tissue Development

Nicolas Willemarck, Evelien Rysman, Koen Brusselmans, Griet Van Imschoot, Frank Vanderhoydonc, Katrien Moerloose, Evelyne Lerut, Guido Verhoeven, Frans van Roy, Kris Vleminckx, and Johannes V. Swinnen

**Précis:** Findings suggest that characteristic elevations in lipogenesis in cancer cells contribute to tumor growth not by improving membrane biosynthesis, as commonly believed, but by modifying fundamental developmental signaling and cell polarity.

9463

Double-Negative Feedback Loop between Reprogramming Factor LIN28 and microRNA *let-7* Regulates Aldehyde Dehydrogenase 1-Positive Cancer Stem Cells

Xiaojun Yang, Xiaojuan Lin, Xiaomin Zhong, Sippy Kaur, Ning Li, Shun Liang, Heini Lassus, Liping Wang, Dionyssios Katsaros, Kathleen Montone, Xia Zhao, Youcheng Zhang, Ralf Bützow, George Coukos, and Lin Zhang

**Précis:** Findings offer evidence that cancer stem cells may arise through cellular reprogramming-like mechanisms.

9473

MicroRNA miR-302 Inhibits the Tumorigenecity of Human Pluripotent Stem Cells by Coordinate Suppression of the CDK2 and CDK4/6 Cell Cycle Pathways

Shi-Lung Lin, Donald C. Chang, Shao-Yao Ying, Davey Leu, and David T.S. Wu

**Précis:** As the most abundant tumor suppressor microRNA in human embryonic stem cells, miR-302 offers an appealing focus for the development of a universal cancer therapeutic.

9483 Silencing of Autocrine Motility Factor Induces Mesenchymal-to-Epithelial Transition and Suppression of Osteosarcoma Pulmonary Metastasis

> Yasufumi Niinaka, Kiyoshi Harada, Masahiro Fujimuro, Masamitsu Oda, Arayo Haga, Misa Hosoki, Narikazu Uzawa, Naoya Arai, Satoshi Yamaguchi, Masashi Yamashiro, and Avraham Raz

**Précis:** An enzyme that functions inside cells in glycolysis and gluconeogenesis has a second extracellular function that in cancer provides a critical support for invasive cell motility, EMT, and metastasis.

9494 The Telomerase Inhibitor Imetelstat
Depletes Cancer Stem Cells in Breast
and Pancreatic Cancer Cell Lines

Immanual Joseph, Robert Tressler, Ekaterina Bassett, Calvin Harley, Christen M. Buseman, Preeti Pattamatta, Woodring E. Wright, Jerry W. Shay, and Ning F. Go

**Précis:** Findings offer preclinical proof-of-concept for a highly tractable and attractive new approach to target cancer stem cells in cancer therapy.

9505 Loss of Thioredoxin Reductase 1 Renders Tumors Highly Susceptible to Pharmacologic Glutathione Deprivation

> Pankaj Kumar Mandal, Manuela Schneider, Pirkko Kölle, Peter Kuhlencordt, Heidi Förster, Heike Beck, Georg W. Bornkamm, and Marcus Conrad

**Précis:** Findings offer preclinical proof-of-concept that simultaneous inhibition of more than one cellular antioxidant system could be highly effective in killing tumor cells via redox modifiers.

9515 Induction of Human Enithelial Stem/Pros

Epithelial Stem/Progenitor Expansion by FOXM1

Emilios Gemenetzidis, Daniela Elena-Costea, Eric K. Parkinson, Ahmad Waseem, Hong Wan, and Muy-Teck Teh

**Précis:** Results identify a transcription factor in the FOXM1 family that can hijack adult human stem cells to jump start cancer initiation.

#### **CORRECTIONS**

9527

9527 Correction: Curcumin Blocks RON
Tyrosine Kinase–Mediated Invasion of

Breast Carcinoma Cells

Correction: RG7204 (PLX4032), a Selective BRAF<sup>V600E</sup> Inhibitor, Displays Potent Antitumor Activity in Preclinical Melanoma Models

#### RETRACTIONS

9529

9528 Retraction: Immunotherapeutic
Potential of B7-DC (PD-L2)
Cross-Linking Antibody In
Conferring Antitumor Immunity

9528 Retraction: An Effective Vaccine Strategy Protective against Antigenically Distinct Tumor Variants

> Retraction: CTL Activation Using the Natural Low-Affinity Epitope 222-229 from Tyrosinase-Related Protein 1

Leads to Tumor Rejection

#### **ABOUT THE COVER**

Selective antibody-mediated,  $\alpha$ -particle–directed killing of LS174T human colon cancer neovascular endothelium using  $^{225}\text{Ac-E4G10}$  "normalizes" vessels. Animals were treated 15 days before sacrifice and tumors were stained for endothelium (green), pericytes (red), and nuclei (blue). This is a representative cross-section of vessels of  $^{225}\text{Ac-E4G10}$  treated tumors. More pericytes were associated with endothelial cells when compared with chemotherapy-treated or saline-treated controls. Boxes are 6- $\mu$ m square. For details, see the article by Escorcia and colleagues on page 9277 of this issue.

