## Highlights of This Issue

### SPECIAL FEATURES

#### CCR Translations

- **Oncolytic Virotherapy Needs Trials, Not Access Programs**
  - Kevin J. Harrington
  - *See article, p. 2734*

#### Molecular Pathways

- **Molecular Pathways: Tumor-Derived Microvesicles and Their Interactions with Immune Cells In Vivo**
  - Ferdinando Pucci and Mikael J. Pittet

#### CCR Focus

- **PFS: The Endpoint We Love and Love to Hate**
  - Susan E. Bates

- **Overview: Progression-Free Survival as an Endpoint in Clinical Trials with Solid Tumors**
  - Ronald L. Korn and John J. Crowley

- **Missing Data and Measurement Variability in Assessing Progression-Free Survival Endpoint in Randomized Clinical Trials**
  - Rajeshwari Sridhara, Sumithra J. Mandrekar, and Lori E. Dodd

- **The Imaging Viewpoint: How Imaging Affects Determination of Progression-Free Survival**
  - Daniel Carl Sullivan, Lawrence H. Schwartz, and Binsheng Zhao

- **The Clinical Viewpoint: Definitions, Limitations of RECIST, Practical Considerations of Measurement**
  - Liza C. Villaruz and Mark A. Socinski

- **Assessment of Audit Methodologies for Bias Evaluation of Tumor Progression in Oncology Clinical Trials**

#### Human Cancer Biology

- **Aberrant BAF57 Signaling Facilitates Prometastatic Phenotypes**
  - Sucharitha Balasubramaniam, Clay E.S. Comstock, Adam Ertel, Kwang Won Jeong, Michael R. Stallcup, Sankar Addya, Peter A. McCue, William F. Ostrander Jr, Michael A. Augello, and Karen E. Knudsen

- **Relapsed Classic E-Cadherin (CDH1)–Mutated Invasive Lobular Breast Cancer Shows a High Frequency of HER2 (ERBB2) Gene Mutations**

#### Cancer Therapy: Preclinical

- **Potent Antimyeloma Activity of a Novel ERK5/CDK Inhibitor**
  - Stela Álvarez-Fernández, Maria Jesús Ortiz-Ruiz, Tracy Parrott, Sara Zaknoen, Enrique M. Ocío, Jesús San Miguel, Francis J. Burrows, Azucena Esparís-Ogando, and Atanasio Pandiella

- **Concomitant BRAF and PI3K/mTOR Blockade Is Required for Effective Treatment of BRAFV600E Colorectal Cancer**
Temozolomide-Mediated DNA Methylation in Human Myeloid Precursor Cells: Differential Involvement of Intrinsic and Extrinsic Apoptotic Pathways

Inhibition of Melanoma Growth by Small Molecules That Promote the Mitochondrial Localization of ATF2

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Antiviral and Antitumor T-cell Immunity in Patients Treated with GM-CSF–Coding Oncolytic Adenovirus
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Sorafenib or Placebo with Either Gemcitabine or Capecitabine in Patients with HER-2–Negative Advanced Breast Cancer That Progressed during or after Bevacizumab

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Matthew C. Winter, Caroline Wilson, Stuart P. Syddall, Simon S. Cross, Alyson Evans, Christine E. Ingram, Ingrid J. Jolley, Matthew Q. Hatton, Jennifer V. Freeman, Stefano Mori, Ingunn Holen, and Robert E. Coleman

Phase I Study of the Hedgehog Pathway Inhibitor IPI-926 in Adult Patients with Solid Tumors

Molecular Profiling of Aromatase Inhibitor–Treated Postmenopausal Breast Tumors Identifies Immune-Related Correlates of Resistance

Correction: Chromosome 5q Loss in Colorectal Flat Adenomas

See commentary, p. 2595
ABOUT THE COVER

This image is taken from a bone metastasis in a patient with stage IV relapsed invasive CDH1 mutated lobular carcinoma of the breast. The tumor was negative for ERBB2 (HER2) amplification (FISH). The targeted next generation sequencing assay used in this study found an ERBB2-GRB7 putative gene fusion that has not been previously reported. The fusion retains the kinase domain of ERBB2 (uniprot.org) which suggests that it could result in ERBB2 activation. The 17q12-21 amplicon which includes both ERBB2 and GRB7 is frequently amplified in breast cancer and preclinical studies suggest that it may be a recombination hotspot. An expression screening study has reported that GRB7 can function as an ERBB2-dependent oncogene. GRB7 encodes an adaptor protein that interacts with ERBB2 and has been shown in a preclinical study to enhance its transformative capacity and increase ERBB2 phosphorylation in fibroblasts. For details, see the article by Ross and colleagues on page 2668 of this issue.

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