## In This Issue

Highlighted research articles .......................... 961

## News in Brief

Important news stories affecting the community ........................ 964

## News in Depth

Q&A: George Sledge on Trends in Clinical Trials ............ 967

Emphasizing the Provocative ............................. 968

## Research Watch

Selected highlights of recent articles of exceptional significance from the cancer literature ............ 969

## Online

For more News and Research Watch, visit Cancer Discovery online at http://CDnews.aacrjournals.org.

## Views

In The Spotlight

The Potential of Circulating Tumor Cells as a Liquid Biopsy to Guide Therapy in Prostate Cancer .................. 974

K. Pantel and C. Alix-Panabières

Commentary on Miyamoto et al., p. 995

Fingerprinting Acute Leukemia: DNA Methylation Profiling of B-Acute Lymphoblastic Leukemia ........... 976

L. Cimmino and I. Aifantis

Commentary on Geng et al., p. 1004

Distinct Epigenetic Mechanisms Distinguishing TMPRSS2-ERG Fusion-Positive and -Negative Prostate Cancers ...................... 979

J.J. Alumkal and J.G. Herman

Commentary on Börno et al., p. 1024

## Research Briefs

### Review

Measuring Oncogenic Signaling Pathways in Cancer with PET: An Emerging Paradigm from Studies in Castration-Resistant Prostate Cancer .......................... 985

M.J. Evans

### Research

Androgen Receptor Signaling in Circulating Tumor Cells as a Marker of Hormonally Responsive Prostate Cancer .................. 995


Précis: Automated immunofluorescence imaging of circulating tumor cells can noninvasively detect androgen receptor activity in patients with metastatic prostate cancer.

### Research Articles

Integrative Epigenomic Analysis Identifies Biomarkers and Therapeutic Targets in Adult B-Acute Lymphoblastic Leukemia ...................... 1004


Précis: Distinct DNA methylation profiles and gene expression patterns are associated with expression of leukemic fusion proteins in adult B-ALLs with poor outcome.
Genome-wide DNA Methylation Events in TMPRSS2–ERG Fusion-Negative Prostate Cancers Implicate an EZH2-Dependent Mechanism with miR-26a Hypermethylation ................. 1024
Précis: EZH2 overexpression is caused by miR-26a hypermethylation in prostate cancers lacking the TMPRSS2–ERG gene fusion, which have distinct DNA methylation profiles.

PI3K Inhibition Impairs BRCA1/2 Expression and Sensitizes BRCA-Proficient Triple-Negative Breast Cancer to PARP Inhibition .............. 1036
Précis: PI3K inhibition synergizes with PARP inhibitors in vivo to decrease the growth of BRCA1-mutant breast tumors, revealing a role for PI3K in the DNA damage response.

Miyamoto and colleagues noninvasively assayed androgen receptor (AR) signaling activity in patients with prostate cancer by measuring levels of prostate-specific antigen (PSA) and prostate-specific membrane antigen (PSMA) in single circulating tumor cells (CTC). The CTCs of untreated patients showed an “AR-on” (PSA+/PSMA−) signature that switched to an “AR-off” (PSA+/PSMA+) signature after androgen deprivation therapy, but the CTCs of patients with castration-resistant prostate cancer (CRPC) were heterogeneous and had “AR-on,” “AR-off,” and “AR-mixed” (PSA+/PSMA+) signatures. The presence of “AR-mixed” CTCs was associated with a poor response to abiraterone acetate, suggesting that monitoring of AR signaling in CTCs may guide use of secondary hormonal therapies in patients with CRPC. For details, please see the article by Miyamoto and colleagues on page 995.