

| | Commentary and Review | |
|---------|---|-------------------------|
| Review | Finding the fifth base: Genome-wide sequencing of cytosine methylation Ryan Lister and Joseph R. Ecker | 959 |
| | Research | |
| Letters | Comparative analysis of H2A.Z nucleosome organization in the human and yeast genomes Michael Y. Tolstorukov, Peter V. Kharchenko, Joseph A. Goldman, Robert E. Kingston, and Peter J. Park | 967 |
| | Large-scale mRNA sequencing determines global regulation of RNA editing during brain development Helene Wahlstedt, Chammiran Daniel, Mats Ensterö, and Marie Öhman | 978 |
| | The colorectal cancer risk at 18q21 is caused by a novel variant altering SMAD7 expression Alan M. Pittman, Silvia Naranjo, Emily Webb, Peter Broderick, Esther H. Lips, Tom van Wezel, Hans Morreau, Kate Sullivan, Sarah Fielding, Philip Twiss, Jayaram Vijayakrishnan, Fernando Casares, Mobshra Qureshi, José Luis Gómez-Skarmeta, and Richard S. Houlston | 987 |
| | Functional diversity for REST (NRSF) is defined by in vivo binding affinity hierarchies at the DNA sequence level Alexander W. Bruce, Andrés J. López-Contreras, Paul Flicek, Thomas A. Down, Pawandeep Dhami, Shane C. Dillon, Christoph M. Koch, Cordelia F. Langford, Ian Dunham, Robert M. Andrews, and David Vetrie | 994^{0A} |
| | Genomic mapping of binding regions for the Ecdysone receptor protein complex Zareen Gauhar, Ling V. Sun, Sujun Hua, Christopher E. Mason, Florian Fuchs, Tong-Ruei Li, Michael Boutros, and Kevin P. White | 1006 |
| | Dynamic reprogramming of transcription factors to and from the subtelomere H. Craig Mak, Lorraine Pillus, and Trey Ideker | 1014 |

| | Comparative inference of illegitimate recombination between rice and sorghum duplicated genes produced by polyploidization Xiyin Wang, Haibao Tang, John E. Bowers, and Andrew H. Paterson | 1026 |
|-----------|---|--------------------|
| | Complete genome of the cellulolytic thermophile Acidothermus cellulolyticus 11B provides insights into its ecophysiological and evolutionary adaptations Ravi D. Barabote, Gary Xie, David H. Leu, Philippe Normand, Anamaria Necsulea, Vincent Daubin, Claudine Médigue, William S. Adney, Xin Clare Xu, Alla Lapidus, Rebecca E. Parales, Chris Detter, Petar Pujic, David Bruce, Celine Lavire, Jean F. Challacombe, Thomas S. Brettin, and Alison M. Berry | 1033 |
| | Methods and Resources | |
| Methods | Distinct DNA methylation patterns characterize differentiated human embryonic stem cells and developing human fetal liver Alayne L. Brunner, David S. Johnson, Si Wan Kim, Anton Valouev, Timothy E. Reddy, Norma F. Neff, Elizabeth Anton, Catherine Medina, Loan Nguyen, Eric Chiao, Chuba B. Oyolu, Gary P. Schroth, Devin M. Absher, Julie C. Baker, and Richard M. Myers | 1044 ^{0a} |
| | Integrating siRNA and protein-protein interaction data to identify an expanded insulin signaling network Zhidong Tu, Carmen Argmann, Kenny K. Wong, Lyndon J. Mitnaul, Stephen Edwards, Iliana C. Sach, Jun Zhu, and Eric E. Schadt | 1057 |
| | High-throughput genotyping by whole-genome resequencing Xuehui Huang, Qi Feng, Qian Qian, Qiang Zhao, Lu Wang, Ahong Wang, Jianping Guan, Danlin Fan, Qijun Weng, Tao Huang, Guojun Dong, Tao Sang, and Bin Han | 1068 ^{0a} |
| | Mapping epigenetic mutations in fission yeast using whole-genome next-generation sequencing Danielle V. Irvine, Derek B. Goto, Matthew W. Vaughn, Yukinobu Nakaseko, W. Richard McCombie, Mitsuhiro Yanagida, and Rob Martienssen | 1077 |
| | Single-pass classification of all noncoding sequences in a bacterial genome using phylogenetic profiles Antonin Marchais, Magali Naville, Chantal Bohn, Philippe Bouloc, and Daniel Gautheret | 1084 |
| Resources | Exploring the human genome with functional maps Curtis Huttenhower, Erin M. Haley, Matthew A. Hibbs, Vanessa Dumeaux, Daniel R. Barrett, Hilary A. Coller, and Olga G. Troyanskaya | 1093 |

| Global networks of functional coupling in eukaryotes from comprehensive data integration | 1107 |
|---|--------------------|
| Andrey Alexeyenko and Erik L.L. Sonnhammer | |
| ABySS: A parallel assembler for short read sequence data Jared T. Simpson, Kim Wong, Shaun D. Jackman, Jacqueline E. Schein, Steven J.M. Jones, and İnanç Birol | 1117 ^{0a} |
| SNP detection for massively parallel whole-genome resequencing Ruiqiang Li, Yingrui Li, Xiaodong Fang, Huanming Yang, Jian Wang, Karsten Kristiansen, and Jun Wang | 1124 |
| High resolution mapping of expression QTLs in heterogeneous stock mice in multiple tissues Guo-Jen Huang, Sagiv Shifman, William Valdar, Martina Johannesson, Binnaz Yalcin, Martin S. Taylor, Jennifer M. Taylor, Richard Mott, and Jonathan Flint | 1133 |

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Cover A distribution of short reads from high-throughput DNA sequencing combined with chromatin immunoprecipitation (ChIP) can reveal positions of nucleosomes on a genome-scale. In this issue, positions of nucleosomes containing the H2A.Z histone variant or H3 tri-methylation at lysine 4 were identified from ChIP-sequencing data. Analysis of the nucleosomal sequences highlights the differences in nucleosome positioning mechanisms between yeast and human and suggests a novel mechanism for organization of H2A.Z nucleosomes. (Cover illustration by Peter Park, Michael Tolstorukov, and Wonyeop Seok. [For details, see Tolstorukov et al., pp. 967–977.])