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Frequent PIK3CA Mutations in Colorectal and Endometrial Tumors With 2 or More Somatic Mutations in Mismatch Repair Genes

S. A. Cohen, E. H. Turner, M. B. Beightol, A. Jacobson, T. A. Gooley, S. J. Salipante, S. Haraldsdottir, C. Smith, S. Scroggins, J. F. Tait, W. M. Grady, E. H. Lin, D. E. Cohn, P. J. Goodfellow, M. W. Arnold, A. de la Chapelle, R. Pearlman, H. Hampel, and C. C. Pritchard

PIK3CA mutations might be used to identify a specific group of colorectal tumors, and to select treatment or determine prognosis.

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Efficacy of the Combination of Sofosbuvir, Velpatasvir, and the NS3/4A Protease Inhibitor GS-9857 in Treatment-Naïve or Previously Treated Patients With Hepatitis C Virus Genotype 1 or 3 Infections

E. J. Gane, C. Schwabe, R. H. Hyland, Y. Yang, E. Svarovskaia, L. M. Stamm, D. M. Brainard, J. G. McHutchison, and C. A. Stedman

Eight weeks of treatment with the combination of sofosbuvir, velpatasvir, and GS-9857 produced a sustained virologic response at 12 weeks after therapy in most treatment-naïve or previously treated patients with HCV genotype 1 or 3.

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G. N. Ioannou, L. A. Beste, M. F. Chang, P. K. Green, E. Lowy, J. I. Tsui, F. Su, and K. Berry

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Sofosbuvir, Ledipasvir/Sofosbuvir, or Paritaprevir/Ritonavir/Ombitasvir and Dasabuvir regimens achieved remarkably high sustained virologic response rates at 12 weeks in patients with HCV genotypes 1–4 infections.

472 Long-term Effects of Hepatitis B Immunization of Infants in Preventing Liver Cancer M.-H. Chang, S.-L. You, C.-J. Chen, C.-J. Liu, M.-W. Lai, T.-C. Wu, S.-F. Wu, C.-M. Lee, S.-S. Yang, H.-C. Chu, T.-E. Wang, B.-W. Chen, W.-L. Chuang, M.-S. Soon, C.-Y. Lin, S.-T. Chiou, H.-S. Kuo, D.-S. Chen, and the Taiwan Hepatoma Study Group

See editorial on page 390.

Immunization of infants against HBV markedly reduced their subsequent risk of developing hepatocellular carcinoma as children and young adults.

Persistent Nonalcoholic Fatty Liver Disease Increases Risk for Carotid Atherosclerosis D. H. Sinn, S. J. Cho, S. Gu, D. Seong, D. Kang, H. Kim, B.-K. Yi, S. W. Paik, E. Guallar, J. Cho, and G.-Y. Gwak

Persistent NAFLD was associated with an increased risk of developing subclinical carotid atherosclerosis. This association was explained by metabolic factors that could potentially mediate the systemic effects of NAFLD.

Basic and Translational—Alimentary Tract

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Commensal-Specific CD4⁺ Cells From Patients With Crohn's Disease Have a T-Helper 17 Inflammatory Profile

E. Calderón-Gómez, H. Bassolas-Molina, R. Mora-Buch, I. Dotti, N. Planell, M. Esteller, M. Gallego, M. Martí, C. Garcia-Martín, C. Martínez-Torró, I. Ordás, S. Singh, J. Panés, D. Benítez-Ribas, and A. Salas

A larger proportion of commensal-specific CD4⁺ T cells from patients with Crohn's disease have a Th17 phenotype and/or produce Th1 and Th17 cytokines, compared with T cells from controls, which might contribute to intestinal inflammation in patients with this disease.

Basic and Translational—Liver

Prevalence of Resistance-Associated Substitutions in HCV NS5A, NS5B, or NS3 and Outcomes of Treatment With Ledipasvir and Sofosbuvir

C. Sarrazin, H. Dvory-Sobol, E. S. Svarovskaia, B. P. Doehle, P. S. Pang, S.-M. Chuang, J. Ma, X. Ding, N. H. Afdhal, K. V. Kowdley, E. J. Gane, E. Lawitz, D. M. Brainard, J. G. McHutchison, M. D. Miller, and H. Mo

Baseline resistance-associated substitutions in the HCV NS5A protein exert minimal effects on patient response to ledipasvir/sofosbuvir therapy, which could be largely overcome by intensifying treatment or extending treatment duration.

Comparison of Gene Expression Patterns Between Mouse Models of Nonalcoholic Fatty Liver

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A. Teufel, T. Itzel, W. Erhart, M. Brosch, X. Y. Wang, Y. O. Kim, W. von Schönfels, A. Herrmann, S. Brückner, F. Stickel, J.-F. Dufour, T. Chavakis, C. Hellerbrand, R. Spang, T. Maass, T. Becker, S. Schreiber, C. Schafmayer, D. Schuppan, and J. Hampe

Little overlap is present in gene expression profiles between liver tissues from different mouse models of NAFLD and patients with different stages of NAFLD.

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526 YAP1 and TAZ Control Pancreatic Cancer Initiation in Mice by Direct Up-regulation

of IAK-STAT3 Signaling

R. Gruber, R. Panayiotou, E. Nye, B. Spencer-Dene, G. Stamp, and A. Behrens

See editorial on page 393.

Oncogenic KRAS facilitates acinar-to-duct metaplasia, thereby generating cells that initiate neoplastic progression. This process involves activation of YAP1 and TAZ in acinar cells, which upregulate JAK -STAT3 signaling to promote development of pancreatic ductal adenocarcinoma in mice.

540 ETS-Transcription Factor ETV1 Regulates Stromal Expansion and Metastasis

in Pancreatic Cancer

S. Heeg, K. K. Das, M. Reichert, B. Bakir, S. Takano, J. Caspers, N. M. Aiello, K. Wu, A. Neesse, A. Maitra, C. A. Iacobuzio-Donahue, P. Hicks, and A. K. Rustgi

ETV1 is critical in the desmoplastic stromal expansion and metastatic progression of pancreatic cancer in mice, and is mediated functionally in part through SPACR2 and HAS2.

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