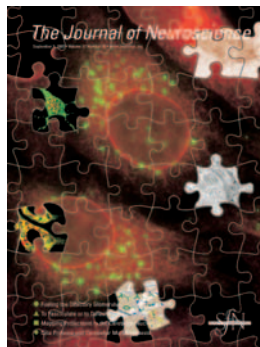


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Cover legend: The cellular effects of the ALS-linked P56S mutation in VAPB. Expression of VAPB-P56S in cultured cells results in formation of endoplasmic reticulum (ER)-derived tubular aggregates (green). A combination of loss of function (disrupted lipid protein binding), dominant-negative effects (wild-type VAP recruitment), and gain of toxic function (disrupted membrane trafficking) may lead to VAPB-linked motor neuron disease. Insets show normal VAPB localization in motor neurons (top right) and its colocalization with ER marker PO (bottom left), immuno-EM of VAPB-P56S aggregates (bottom right), Golgi fragmentation in VAPB-P56S-expressing neurons (top left), and the VAPA protein structure (bottom middle). The cover was designed by Eva Teuling and Casper Hoogenraad. For more information, see the article by Teuling et al. in this issue (pages 9801–9815).

i This Week in The Journal

Journal Club

- 9535 **Labeled-Line Coding and Summation Coding of Numerosities in Prefrontal and Parietal Cortex**
Filip Van Opstal

Brief Communications

- 9648 **Sequential Development of Long-Term Potentiation and Depression in Different Layers of the Mouse Visual Cortex**
Bin Jiang, Mario Treviño, and Alfredo Kirkwood
- 9664 **Nuclear Factor- κ B Activation via Tyrosine Phosphorylation of Inhibitor κ B- α Is Crucial for Ciliary Neurotrophic Factor-Promoted Neurite Growth from Developing Neurons**
Denis Gallagher, Humberto Gutierrez, Nuria Gavaldà, Gerard O’Keeffe, Ron Hay, and Alun M. Davies

Articles

CELLULAR/MOLECULAR

- 9560 **Synaptic Kainate Receptors Tune Oriens-Lacunosum Moleculare Interneurons to Operate at Theta Frequency**
Miri Goldin, Jérôme Epszstein, Isabel Jorquera, Alfonso Represa, Yehzekel Ben-Ari, Valérie Crépel, and Rosa Cossart
- 9573 **Axonal Neurofilaments Control Multiple Fiber Properties But Do Not Influence Structure or Spacing of Nodes of Ranvier**
Rodolphe Perrot, Pierre Lonchampt, Alan C. Peterson, and Joël Eyer
- 9711 **Spike-Timing-Dependent Plasticity of Neocortical Excitatory Synapses on Inhibitory Interneurons Depends on Target Cell Type**
Jiang-teng Lu, Cheng-yu Li, Jian-Ping Zhao, Mu-ming Poo, and Xiao-hui Zhang
- 9721 **Compartment-Specific Modulation of GABAergic Synaptic Transmission by μ -Opioid Receptor in the Mouse Striatum with Green Fluorescent Protein-Expressing Dopamine Islands**
Masami Miura, Sachiko Saino-Saito, Masao Masuda, Kazuto Kobayashi, and Toshihiko Aosaki
- 9736 **Extracellular Glutamate Concentration in Hippocampal Slice**
Melissa A. Herman and Craig E. Jahr

- 9790 **An Energy Budget for the Olfactory Glomerulus**
Janna C. Nawroth, Charles A. Greer, Wei R. Chen, Simon B. Laughlin, and
Gordon M. Shepherd

DEVELOPMENT/PLASTICITY/REPAIR

- 9653 **Semaphorin3D Regulates Axon–Axon Interactions by Modulating Levels of L1 Cell Adhesion Molecule**
Marc A. Wolman, Ann M. Regnery, Thomas Becker, Catherina G. Becker, and
Mary C. Halloran
- 9670 **Fate-Mapping the Mammalian Hindbrain: Segmental Origins of Vestibular Projection Neurons Assessed Using Rhombomere-Specific *Hoxa2* Enhancer Elements in the Mouse Embryo**
Massimo Pasqualetti, Carmen Díaz, Jean-Sébastien Renaud, Filippo M. Rijli, and
Joel C. Glover
- 9682 **Delineation of Multiple Subpallial Progenitor Domains by the Combinatorial Expression of Transcriptional Codes**
Nuria Flames, Ramón Pla, Diego M. Gelman, John L. R. Rubenstein, Luis Puelles,
and Oscar Marín
- 9757 **Activation of the *Wnt*– β Catenin Pathway in a Cell Population on the Surface of the Forebrain Is Essential for the Establishment of Olfactory Axon Connections**
Ambra A. Zaghetto, Sara Paina, Stefano Mantero, Natalia Platonova,
Paolo Peretto, Serena Bovetti, Adam Puche, Stefano Piccolo, and Giorgio R. Merlo

BEHAVIORAL/SYSTEMS/COGNITIVE

- 9585 **Human Adult Cortical Reorganization and Consequent Visual Distortion**
Daniel D. Dilks, John T. Serences, Benjamin J. Rosenau, Steven Yantis, and
Michael McCloskey
- 9607 **State Changes Rapidly Modulate Cortical Neuronal Responsiveness**
Andrea Hasenstaub, Robert N. S. Sachdev, and David A. McCormick
- 9623 **Responses of Suprachiasmatic Nucleus Neurons to Light and Dark Adaptation: Relative Contributions of Melanopsin and Rod–Cone Inputs**
Elise Drouyer, Camille Rieux, Roelof A. Hut, and Howard M. Cooper
- 9632 **Spatial Attention and the Latency of Neuronal Responses in Macaque Area V4**
Joonyeol Lee, Tori Williford, and John H. R. Maunsell
- 9638 **Computational Diversity in Complex Cells of Cat Primary Visual Cortex**
Ian M. Finn and David Ferster
- 9696 **Molecular, Topographic, and Functional Organization of the Cerebellar Nuclei: Analysis by Three-Dimensional Mapping of the Olivonuclear Projection and Aldolase C Labeling**
Izumi Sugihara and Yoshikazu Shinoda
- 9729 **Estrogen Disrupts the Inhibition of Fear in Female Rats, Possibly through the Antagonistic Effects of Estrogen Receptor α (ER α) and ER β**
Donna J. Toufexis, Karyn M. Myers, Michael E. Bowser, and Michael Davis
- 9742 **Multimodal Coding of Three-Dimensional Rotation and Translation in Area MSTd: Comparison of Visual and Vestibular Selectivity**
Katsumasa Takahashi, Yong Gu, Paul J. May, Shawn D. Newlands,
Gregory C. DeAngelis, and Dora E. Angelaki
- 9769 **Hippocampal CA1 Place Cells Encode Intended Destination on a Maze with Multiple Choice Points**
James A. Ainge, Minija Tamosiunaite, Florentin Woergoetter, and
Paul A. Dudchenko

NEUROBIOLOGY OF DISEASE

- 9537 **Nonpsychoactive Cannabidiol Prevents Prion Accumulation and Protects Neurons against Prion Toxicity**
Sevda Dirikoc, Suzette A. Priola, Mathieu Marella, Nicole Zsürger, and Joëlle Chabry
- 9545 **Skin-Derived Precursors Generate Myelinating Schwann Cells That Promote Remyelination and Functional Recovery after Contusion Spinal Cord Injury**
Jeff Biernaskie, Joseph S. Sparling, Jie Liu, Casey P. Shannon, Jason R. Plemel, Yuanyun Xie, Freda D. Miller, and Wolfram Tetzlaff
- 9595 **Noradrenergic Modulation of Subthalamic Nucleus Activity: Behavioral and Electrophysiological Evidence in Intact and 6-Hydroxydopamine-Lesioned Rats**
Pauline Belujon, Erwan Bezar, Anne Taupignon, Bernard Bioulac, and Abdelhamid Benazzouz
- 9780 **Cilia Proteins Control Cerebellar Morphogenesis by Promoting Expansion of the Granule Progenitor Pool**
Victor V. Chizhikov, James Davenport, Qihong Zhang, Evelyn Kim Shih, Olga A. Cabello, Jannon L. Fuchs, Bradley K. Yoder, and Kathleen J. Millen
- 9801 **Motor Neuron Disease-Associated Mutant Vesicle-Associated Membrane Protein-Associated Protein (VAP) B Recruits Wild-Type VAPs into Endoplasmic Reticulum-Derived Tubular Aggregates**
Eva Teuling, Suaad Ahmed, Elize Haasdijk, Jeroen Demmers, Michel O. Steinmetz, Anna Akhmanova, Dick Jaarsma, and Casper C. Hoogenraad