

VOLUME 275  
DECEMBER 1995  
NUMBER 3

# The Journal of PHARMACOLOGY AND EXPERIMENTAL THERAPEUTICS

## Contents

### ANALGESIA AND DRUGS OF ABUSE

**Striatal Dopamine Receptors and Adenylyl Cyclase Activity in a Rat Model of Alcohol Addiction: Effects of Ethanol and Lusiride Treatment**

**Regulation of Morphine Antiallodynic Efficacy by Cholecystokinin in a Model of Neuropathic Pain in Rats**

**Antinociceptive Effects of Cocaine/Opioid Combinations in Rhesus Monkeys**

**Opioid Antinociception in a Rat Model of Visceral Pain: Systemic Versus Local Drug Administration**

**Participation of  $\Delta$  Opioid Receptor Subtypes in the Stimulation of Adenylyl Cyclase Activity in Rat Olfactory Bulb**

**In Vivo Injection of Antisense Oligodeoxynucleotides to  $G\alpha$  Subunits and Supraspinal Analgesia Evoked by Mu and Delta Opioid Agonists**

### AUTONOMIC PHARMACOLOGY

**Endothelin Subtypes: Effect on Isolated Rhesus Monkey Ciliary Muscle**

**Intervessel (Arteries and Veins) and Heart/Vessel Selectivities of Therapeutically Used Calcium Entry Blockers: Variable, Vessel-Dependent Indexes**

**Pharmacological Evaluation of 1229U91, A Novel High-Affinity and Selective Neuropeptide Y-Y<sub>1</sub> Receptor Antagonist**

**The Expression of Alpha<sub>1</sub> Adrenoceptor Subtypes Changes with Age in the Rat Aorta**

### BEHAVIORAL PHARMACOLOGY

**Effects of Chronic Haloperidol on Reaction Time and Errors in a Sustained Attention Task: Partial Reversal by Anticholinergics and by Amphetamine**

**Nafadotride, a Potent Preferential Dopamine D<sub>3</sub> Receptor Antagonist, Activates Locomotion in Rodents**

**Discriminative Stimulus Effects of R-(+)-3-Amino-1-Hydroxypyrrolid-2-One, [(+)-HA-966], a Partial Agonist of the Strychnine-Insensitive Modulatory Site of the N-Methyl-D-Aspartate Receptor**

**The Reinforcing Effects of Dopamine D<sub>1</sub> Receptor Agonists in Rhesus Monkeys**

**Serotonergic Modulation of the Behavioral Effects of Cocaine in the Squirrel Monkey**

Torsten May, Ulrike Wolf and Jochen Wolffgramm 1195

Michael L. Nichols, Di Bian, Michael H. Ossipov, Josephine Lai and Frank Porreca 1339

Michael B. Gatch, S. Stevens Negus, Eduardo R. Butelman and Nancy K. Mello 1346

R. M. Craft, S. R. Henley, R. C. Haaseth, V. J. Hruby and F. Porreca 1535

Maria C. Olianas and Pierluigi Onali 1560

Pilar Sánchez-Blázquez, Antonio García-España and Javier Garzón 1590

Cameron Millar, John F. Poyer, B'Ann T. Gabelt and Paul L. Kaufman 1143

Martine Magnon, Pierre Gallix and Icilio Cavero 1157

S. S. Hegde, D. W. Bonhaus, W. Stanley, R. M. Eglen, T. M. Moy, M. Loeb, S. G. Shetty, A. DeSouza and J. Krstenansky 1261

Hakan Gurdal, Nanda Tilakaratne, R. Dale Brown, Maria Fonseca, Eitan Friedman and Mark D. Johnson 1656

Becky J. Brockel and Stephen C. Fowler 1090

Francois Sautel, Nathalie Griffon, Pierre Sokoloff, Jean-Charles Schwartz, Corinne Launay, Philippe Simon, Jean Costentin, Angele Schoenfelder, Fabrice Garrido, Andre Mann and Camille G. Wermuth 1239

Jeffrey M. Witkin, Scott Brave, Dawn French and Beth Geter-Douglass 1267

Michael R. Weed and William L. Woolverton 1367

Leonard L. Howell and Larry D. Byrd 1551

**CARDIOVASCULAR PHARMACOLOGY**

<b>Diabetes and ATP-Sensitive Potassium Channel Openers and Blockers in Isolated Ischemic/Reperfused Hearts</b>	Arpad Tosaki, Daniel T. Engelman, Richard M. Engelman and Dipak K. Das	1115
<b>Effect of a New Aminopeptidase P Inhibitor, Apstatin, on Bradykinin Degradation in the Rat Lung</b>	M. Margaret Prechel, Arthur T. Orawski, Linda L. Maggiora and William H. Simmons	1136
<b>Ischemic Preconditioning, Adenosine and Bethanechol Protect Spontaneously Hypertensive Isolated Rat Hearts</b>	Ashraf Boutros and Jun Wang	1148
<b>1,3-Dipropyl-8-[2-(5,6-Epoxy)Norbornyl]Xanthine, a Potent, Specific and Selective A<sub>1</sub> Adenosine Receptor Antagonist in the Guinea Pig Heart and Brain and in DDT<sub>1</sub>MF-2 Cells</b>	Luiz Belardinelli, John C. Shryock, Yi Zhang, Peter J. Scammells, Ray Olsson, Donn Dennis, Peter Milner, Jürg Pfister and Stephen P. Baker	1167
<b>Suppression by Mycophenolate Mofetil of the Neointimal Thickening Caused by Vascular Injury in a Rat Arterial Stenosis Model</b>	Elizabeth B. Fraser-Smith, Jose D. Rosete and Randall C. Schatzman	1204
<b>Hypoxic Relaxation in Functionally Intact Cattle Coronary Artery Segments Involves K<sup>+</sup> ATP Channels</b>	Stanley Kalsner	1219
<b>Vasorelaxing Effect of S-Nitrosocaptopril on Dog Coronary Arteries: No Cross-tolerance with Nitroglycerin</b>	Tetsuya Matsumoto, Masayuki Takahashi, Ichiro Nakae and Masahiko Kinoshita	1247
<b>Growth Inhibitory Effects on Human Colon Adenocarcinoma-Derived Caco-2 Cells and Calcemic Potential of 1<math>\alpha</math>,25-Dihydroxyvitamin D<sub>3</sub> Analogs: Structure-Function Relationships</b>	M. G. Bischof, K. Redlich, C. Schiller, M. V. Chirayath, M. Uskoković, M. Peterlik and H. S. Cross	1254
<b>Hemodynamic Effects of Acute and Chronic Treatment with Aladotril, a Mixed Inhibitor of Neutral Endopeptidase and Angiotensin I-Converting Enzyme, in Conscious Rats with Myocardial Infarction</b>	Christine Marie, Claude Mossiat, Jeanne-Marie Lecomte, Jean-Charles Schwartz and Jean Bralet	1324
<b>Short-Term Drug Effects on the Signal-Averaged Electrocardiogram in Healthy Men: Assessment of Intra- and Interindividual Variability of Spectral Temporal Mapping and Time-Domain Analysis</b>	Michael Wolzt, Leopold Schmetterer, Johannes Kastner, Kurt Krejcy, Gabriele Zanaschka, Christian Unfried and Hans-Georg Eichler	1375
<b>Comparison Between the Effects of a Novel Ca<sup>++</sup> Sensitizer and a Phosphodiesterase Inhibitor on Stunned Myocardium</b>	B. Korbmacher, U. Sunderdiek, H. D. Schulte, G. Arnold and J. D. Schipke	1433
<b>Thromboxane Synthase Inhibitors Suppress More Effectively the Aggregation of Thromboxane Receptor-desensitized Than That of Normal Platelets: Role of Adenylylcyclase Up-regulation</b>	Roberta Vezza, Giuseppe G. Nenci and Paolo Gresele	1497
<b>Pharmacology of L-754,142, a Highly Potent, Orally Active, Nonpeptidyl Endothelin Antagonist</b>	D. L. Williams, Jr., K. L. Murphy, N. A. Nolan, J. A. O'Brien, D. J. Pettibone, S. D. Kivlighn, S. M. Krause, E. V. Lis, Jr., G. J. Zingaro, R. A. Gabel, F. C. Clayton, P. K. S. Siegl, K. Zhang, J. Naue, K. Vyas, T. F. Walsh, K. J. Fitch, P. K. Chakravarty, W. J. Greenlee and B. V. Clineschmidt	1518
<b>The Specific Contribution of the Novel Alpha-1D Adrenoceptor to the Contraction of Vascular Smooth Muscle</b>	Michael T. Piascik, Richard D. Guarino, Marta S. Smith, Edward E. Soltis, David L. Saussy, Jr. and Dianne M. Perez	1583

**CELLULAR AND MOLECULAR PHARMACOLOGY**

<b>The Caffeine- and Ryanodine-Sensitive Ca<sup>++</sup> Store in Porcine Myometrial Cells: Its Heterogeneity of All-or-None Ca<sup>++</sup> Release</b>	Ronghua Zhuge and Walter H. Hsu	1077
<b>Endothelin ET<sub>A</sub> and ET<sub>B</sub> Receptors Facilitating Parasympathetic Neurotransmission in the Rabbit Trachea</b>	Tomoko Yoneyama, Masatoshi Hori, Takeo Tanaka, Yuzuru Matsuda and Hideaki Karaki	1084
<b>Pharmacology of a Constitutively Active Muscarinic Receptor Generated by Random Mutagenesis</b>	Tracy A. Spalding, Ethan S. Burstein, Hans Brauner-Osborne, David Hill-Eubanks and Mark R. Brann	1274

<b>Cholecystokinin Octapeptide Reverses <math>\mu</math>-Opioid-Receptor-Mediated Inhibition of Calcium Current in Rat Dorsal Root Ganglion Neurons</b>	Nai-Jiang Liu, Tao Xu, Chang Xu, Chong-Qing Li, Ying-Xin Yu, Hua-Guang Kang and Ji-Sheng Han	1293
<b>Methylmercury Acts at Multiple Sites to Block Hippocampal Synaptic Transmission</b>	Yukun Yuan and William D. Atchison	1308
<b>Selective Block of Tetramethrin-Modified Sodium Channels by (<math>\pm</math>)-<math>\alpha</math>-Tocopherol (Vitamin E)</b>	Jin-Ho Song and Toshio Narahashi	1402
<b>The "Calcium Antagonist" TMB-8 [3,4,5-Trimethoxybenzoic acid 8-(Diethylamino)Octyl Ester] Is a Potent, Non-competitive, Functional Antagonist at Diverse Nicotinic Acetylcholine Receptor Subtypes</b>	Merouane Bencherif, Cynthia M. Eisenhour, Richard J. Prince, Patrick M. Lippiello and Ronald J. Lukas	1418
<b>Tenidap, an Anti-inflammatory Agent, Discharges Intracellular <math>\text{Ca}^{++}</math> Store and Inhibits <math>\text{Ca}^{++}</math> Influx in Cultured Human Gingival Fibroblasts</b>	Akira Fujii, Hiroko Matsumoto, Takafumi Hashimoto and Yoshiaki Akimoto	1447
<b>Differential Effects of 2,4-Dithiobiuret on the Synthesis and Release of Acetylcholine and Dopamine from Rat Pheochromocytoma (PC12) Cells</b>	Lynne M. Ireland, Chun-Hong Yan, Laura M. Nelson and William D. Atchison	1453
<b>Potential Risk of Myopathy by HMG-CoA Reductase Inhibitors: A Comparison of Pravastatin and Simvastatin Effects on Membrane Electrical Properties of Rat Skeletal Muscle Fibers</b>	Sabata Pierno, Annamaria De Luca, Domenico Tricarico, Ada Roselli, Fedele Natuzzi, Ermanno Ferrannini, Marcella Laico and Diana Conte Camerino	1490
<b>Angiotensin II Tachyphylaxis in the Guinea Pig Ileum and its Prevention: A Pharmacological and Biochemical Study</b>	Celia A. Kanashiro, Therezinha B. Paiva, Antonio C. M. Paiva, Rogerio N. Prioste, Jeannine Aboulafia and Suma I. Shimuta	1543
<b>Cloning and Pharmacological Characterization of the Rabbit Bradykinin B<sub>2</sub> Receptor</b>	Dimcho R. Bachvarov, Elise Saint-Jacques, Jean-François Larrivée, Luc Levesque, Francis Rioux, Guy Drapeau and François Marceau	1623
<b>Long-Term Stimulation of Nicotinic Receptors is Required to Increase Proenkephalin A mRNA Levels and the Delayed Secretion of [<math>\text{Met}^5</math>]-Enkephalin in Bovine Adrenal Medullary Chromaffin Cells</b>	Hong W. Suh, Pearlie M. Hudson, Michael K. McMillian, Kaberi P. Das, Belinda C. Wilson, Gen C. Wu and Jau S. Hong	1663
<b>CHEMOTHERAPY/GENE THERAPY</b>		
<b>Accumulation of Protein-Coated Liposomes in an Extravascular Site: Influence of Increasing Carrier Circulation Lifetimes</b>	Shane A. Longman, Paul G. Tardi, Mike J. Parr, Lewis Choi, Pieter R. Cullis and Marcel B. Bally	1177
<b>Distribution of 1,3-bis(2-Chloroethyl)-1-Nitrosourea and Tracers in the Rabbit Brain After Interstitial Delivery by Biodegradable Polymer Implants</b>	J. F. Strasser, L. K. Fung, S. Eller, S. A. Grossman and W. M. Saltzman	1647
<b>Metallothionein Expression and Resistance to Cisplatin in a Human Germ Cell Tumor Cell Line</b>	Jim Koropatnick, Debra M. Kloth, Salam Kadhim, Joseph L. Chin and M. George Cherian	1681
<b>DEVELOPMENTAL PHARMACOLOGY</b>		
<b>Development of <math>\Delta</math>-Opioid Receptor Subtypes and the Regulatory Role of Weaning: Radioligand Binding Autoradiography and <i>In Situ</i> Hybridization Studies</b>	I. Kitchen, F. M. Leslie, M. Kelly, R. Barnes, T. J. Crook, R. G. Hill, A. Borsodi, G. Toth, P. Melchiorri and L. Negri	1597
<b>DRUG METABOLISM AND DISPOSITION</b>		
<b>Effect of Fluoxetine, Norfluoxetine, Sertraline and Desmethyl Sertraline on Human CYP3A Catalyzed 1'-Hydroxy Midazolam Formation <i>In Vitro</i></b>	Barbara J. Ring, Shelly N. Binkley, Lorin Roskos and Steven A. Wrighton	1131
<b>The Metabolic Formation of Reactive Intermediates from Clozapine, a Drug Associated with Agranulocytosis in Man</b>	J. L. Maggs, D. Williams, M. Pirmohamed and B. K. Park	1463
<b>Clozapine is Oxidized by Activated Human Neutrophils to a Reactive Nitrenium Ion that Irreversibly Binds to the Cells</b>	Zhao Chao Liu and Jack P. Uetrecht	1476
<b>Acetaminophen Does Not Decrease Hepatic 3'-Phosphoadenosine 5'-Phosphosulfate in Mice</b>	Hyo J. Kim, Peter Rozman and Curtis D. Klaassen	1506

<b>Potent and Selective Inactivation of Human Liver Microsomal Cytochrome P-450 Isoforms by L-754,394, an Investigational Human Immune Deficiency Virus Protease Inhibitor</b>	Masato Chiba, Joy A. Nishime and Jiunn H. Lin	1527
<b>Inhibition of the Rat Cytochrome P450 3A2 by an Antisense Phosphorothioate Oligodeoxynucleotide <i>In Vivo</i></b>	John P. Desjardins and Patrick L. Iversen	1608
<b>GASTROINTESTINAL PHARMACOLOGY</b>		
<b>H<sub>3</sub>-Receptor Activation Inhibits Cholinergic Stimulation of Acid Secretion in Isolated Rabbit Fundic Glands</b>	André Bado, Jean-Pierre Laigneau, Laurent Moizo, Yacine Cherifi and Miguel J. M. Lewin	1099
<b>Pharmacological Role and Degradation Processes of Neuromedin N in the Gastrointestinal Tract: An <i>In Vitro</i> and <i>In Vivo</i> Study</b>	H. Barelli, Z. Woskowska, S. Cipris, J. E. T. Fox-Threlkeld, E. E. Daniel, J. P. Vincent and F. Checler	1300
<b>Intestinal Absorption of Calcium <i>In Vivo</i> is Dependent on Endogenous Nitric Oxide</b>	R. Schleifer, M. Galluser and F. Raul	1427
<b>Cloning and Characterization of a Rat H<sup>+</sup>/Peptide Cotransporter Mediating Absorption of <math>\beta</math>-Lactam Antibiotics in the Intestine and Kidney</b>	Hideyuki Saito, Masahiro Okuda, Tomohiro Terada, Sei Sasaki and Ken-Ichi Inui	1631
<b>IMMUNOPHARMACOLOGY</b>		
<b>Mouse Strain Differences in Immunosuppression by Opioids <i>In Vitro</i></b>	Toby K. Eisenstein, Joseph J. Meissler, Jr., Thomas J. Rogers, Ellen B. Geller and Martin W. Adler	1484
<b>NEUROPHARMACOLOGY</b>		
<b>Methamphetamine-Induced Hyperthermia and Dopaminergic Neurotoxicity in Mice: Pharmacological Profile of Protective and Nonprotective Agents</b>	David S. Albers and Patricia K. Sonsalla	1104
<b>Relative Vulnerability of Dopamine and GABA Neurons in Mesencephalic Culture to Inhibition of Succinate Dehydrogenase by Malonate and 3-Nitropropionic Acid and Protection by NMDA Receptor Blockade</b>	Gail D. Zeevalk, Ethel Derr-Yellin and William J. Nicklas	1124
<b>Pharmacodynamic Model for Acute Tolerance Development to the Electroencephalographic Effects of Alfentanil in the Rat</b>	Jaap W. Mandema and D. Russell Wada	1185
<b>Pharmacology of 5-Chloro-7-Trifluoromethyl-1,4-Dihydro-2,3-Quinoxalinedione: A Novel Systemically Active Ionotropic Glutamate Receptor Antagonist</b>	R. M. Woodward, J. E. Huettner, M. Tran, J. Guastella, J. F. W. Keana and E. Weber	1209
<b>Neurochemical and Functional Characterization of the Preferentially Selective Dopamine D3 Agonist PD 128907</b>	T. A. Pugsley, M. D. Davis, H. C. Akunne, R. G. MacKenzie, Y. H. Shih, G. Damsma, H. Wikstrom, S. Z. Whetzel, L. M. Georgic, L. W. Cooke, S. B. DeMattos, A. E. Corbin, S. A. Glase, L. D. Wise, D. Dijkstra and T. G. Heffner	1355
<b>BIII 277 CL Is a Potent and Specific Ion-channel Blocker of the NMDA Receptor-Channel Complex</b>	Adrian J. Carter, Wolf Dietrich Bechtel, Matthias Grauert, Paul Harrison, Herbert Merz and Werner Stransky	1382
<b>U-89843A is a Novel Allosteric Modulator of <math>\gamma</math>-Aminobutyric Acid<sub>A</sub> Receptors</b>	Haesook K. Im, Wha Bin Im, Jeff F. Pregenzer, Don B. Carter and Beverly J. Hamilton	1390
<b>Influence of the Estrus Cycle on the Discrimination of Apparent Neuroactive Steroid Site Subtypes on the <math>\gamma</math>-Aminobutyric Acid<sub>A</sub> Receptor Complex in the Rat</b>	Linda D. McCauley and Kelvin W. Gee	1412
<b>Effects of Chronic Pentobarbital Treatment on the GABA<sub>A</sub> Receptor Complex in Mammalian Cortical Neurons</b>	Rong Yu and Maharaj K. Ticku	1442
<b>Effects of Propofol on Nociceptive Response and Power Spectra of Electroencephalographic and Systemic Arterial Pressure Signals in the Rat: Correlation with Plasma Concentration</b>	Chen-Hsien Yang, Ming-Hwang Shyr, Terry B. J. Kuo, Peter P. C. Tan and Samuel H. H. Chan	1568

<b>5-Hydroxytryptamine<sub>2A</sub> (5-HT<sub>2A</sub>) Receptor Desensitization can Occur Without Down-Regulation</b>	Bryan L. Roth, Esa-Pekka Palvimaki, Sally Berry, Naseem Khan, Nancy Sachs, Ahmet Uluer and M. Salmon Choudhary	1638
<b>Pharmacological Studies of the Regulation of Chronic FOS-Related Antigen Induction by Cocaine in the Striatum and Nucleus Accumbens</b>	Heather E. Nye, Bruce T. Hope, Max B. Kelz, Michael Iadarola and Eric J. Nestler	1671
<b>PROSTAGLANDINS, LEUKOTRIENES AND OTHER EICOSANOIDS</b>		
<b>Prostaglandins Mediate the Stimulatory Effects of Endothelin-1 on cAMP Accumulation and Inositol-1,4,5-trisphosphate Production and Contraction in Cat Iris Sphincter</b>	Sardar Y. K. Yousufzai, Zhi Ye and Ata A. Abdel-Latif	1280
<b>Characterization of 5-Lipoxygenase Inhibitors in Biochemical and Functional <i>In Vivo</i> Assays</b>	Walter G. Smith, Alex F. Shaffer, Jerry L. Currie, Janice M. Thompson, Suzanne Kim, Tadimeti Rao and Peter C. Isakson	1332
<b>Stimulatory Effect of Leminoprazole on Secretion and Synthesis of Mucus by Rabbit Gastric Mucosal Cells</b>	Satoru Takahashi, Eiji Nakamura and Susumu Okabe	1396
<b>PULMONARY PHARMACOLOGY</b>		
<b>Pharmacological Evaluation of Selected, Orally Active, Peptidyl Inhibitors of Human Neutrophil Elastase</b>	M. J. Janusz, S. L. Durham, C. M. Hare, J. L. Geary, A. K. Mandagere, J. C. Poole, T. N. Thompson, D. Xu, M. R. Angelastro, J. P. Burkhardt, T. M. Chen, A. L. Marquart, N. P. Peet and K. K. Hwang	1233
<b>Stimulation of Opioid Mu-Receptors Potentiates Beta Adrenoceptor-Mediated Relaxation of Canine Airway Smooth Muscle</b>	E. Tagaya, J. Tamaoki, A. Chiyo and K. Konno	1288
<b>RENAL PHARMACOLOGY</b>		
<b>L(+) and D(-)-Lactate Modulate Rat Renal Tubular Accumulation of Amantadine in the Presence and Absence of Bicarbonate</b>	Miguel R. Escobar, Kerry Goralski and Daniel S. Sitar	1317
<b>TOXICOLOGY</b>		
<b>A Metalloporphyrin Superoxide Dismutase Mimetic Protects Against Paraquat-Induced Endothelial Cell Injury, <i>In Vitro</i></b>	Brian J. Day, Scott Shawen, Stefan I. Liochev and James D. Crapo	1227
<b>Role of Kupffer Cells in Reperfusion Injury in Fat-Loaded Livers from Ethanol-Treated Rats</b>	Zhi Zhong, Henry D. Connor, Ronald P. Mason, Wei Qu, Wenshi Gao, John J. Lemasters and Ronald G. Thurman	1512
<b>Evidence for Dissimilar Mechanisms of Enhancement of Inorganic and Organic Hydroperoxide Cytotoxicity by L-Histidine</b>	Andrea Guidarelli, Piero Sestili, Andrea Cossarizza, Claudio Franceschi, Flaminio Cattabeni and Orazio Cantoni	1575
<b>Inactivation and Degradation of Human Cytochrome P4502E1 by CCl<sub>4</sub> in a Transfected HepG2 Cell Line</b>	Yan Dai and Arthur I. Cederbaum	1614
<b>Index, Volume 275, October–December, 1995</b>		1688