BioMed. Chem. **1994**, *2*, 743

Development of Dual-Acting Agents for Thromboxane Receptor Antagonism and Thromboxane Synthase Inhibition. 1. Synthesis, Structure Activity relationship, and Evaluation of Substituted ω-Phenyl-ω-(3-Pyridyl)alkenoic Acids.

K. Takeuchi, A.M. Happ, D. E. Mais, N. Layman, B.G. Utterback, V. L. Wyss, J. A. Jakubowski
Lilly Research Laboratories, Eli Lilly and Company, Indianapolis, Indiana 46285

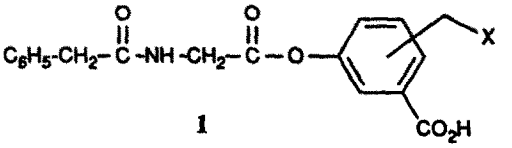
A series of arylsulfonamido-substituted ω-Phenyl-ω-(3-Pyridyl)alkenoic Acids were synthesized and evaluated in vitro for their ability to act as both a thromboxane A_2 receptor antagonist and thromboxane synthase inhibitor.

BioMed. Chem. **1994**, *2*, 757

FUNCTIONALIZED DEPSIPEPTIDES, SUBSTRATES AND INHIBITORS OF β-LACTAMASES AND D,D-PEPTIDASES,

D. Cabaret,[a] J. Liu,[a] M. Wakselman,*[a] R.F. Pratt,*[b] and Y. Xu,[b] *[a]CNRS-CERCOA, 2 rue Henri Dunant, F-94320 Thiais, France and [b]Department of Chemistry, Wesleyan University, Middletown, Connecticut 06459, USA.*

Abstract: Aryl phenaceturates **1** possessing a latent *o*- or *p*-quinone methide function are effective substrates and inhibitors of serine β-lactamases.



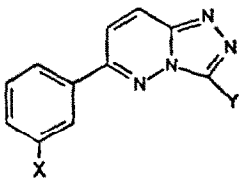
1

X= OAc, $OC(O)C_6H_3(CF_3)_2$, Br, Cl, $S^+(Et)Me\ BF_4^-$.

BioMed. Chem. **1994**, *2*, 773

Bicyclic [*b*]-Heteroannulated Pyridazine Derivatives. 2. Structure-Activity Relationships in the 6-Aryltriazolo[4,3-*b*]pyridazine Ligands of the Benzodiazepine Receptor

J.Karolak-Wojciechowska, J.Lange, W.Kwiatkowski, M.Gniewosz, J.Plenkiewicz
Institute of General and Ecological Chemistry, Technical University, 90924 Łódź, Poland
Chemistry Department, University of Technology, 00662 Warsaw, Poland



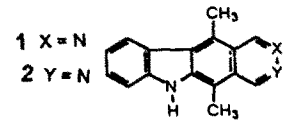
Some electronic parameters have been calculated by semiempirical quantum chemistry methods for two series of the title ligands. The receptor affinity of the compounds correlated well with the ionization potential values. Dipole moment orientation was considered to be another important parameter controlling the ligand-to-receptor binding.

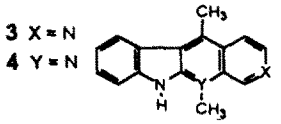
BioMed. Chem. **1994**, *2*, 781

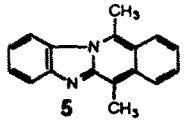
Antineoplastic Activity of Benzimidazo[1,2-b]isoquinolines, Indolo[2,3-b]quinolines, and Pyridocarbazoles

Ronni L. Weinkauf[1], Allan Y. Chen[2], Chiang Yu[2], Leroy Liu[2], Louis Barrows[3], and Edmond J. LaVoie[1]*
[1]Dept. of Pharm. Chem., Rutgers University, Piscataway, N.J. 08855, [2]Dept. of Pharmacol., UMDNJ-R.WJ-Medical School, Piscataway, N.J. 08855, [3]Dept. of Pharmacol. and Toxicol, The University of Utah, Salt Lake City, Utah 84112

The SAR of heterocycles related to ellipticine (**1**) was determined. Compounds **2-5** and analogs of **5** were synthesied and evaluated as mammalian topoisomerase II inhibitors and for cytotoxicity in human tumor cells.







5

BioMed. Chem. **1994**, *2*, 787

SYNTHESIS OF AN IMMUNOLOGICALLY ACTIVE ANALOG OF THYMIC HUMORAL FACTOR-γ2 ENHANCED ENZYMATIC STABILITY, T. Abiko* and H. Sekino, Kidney Research Laboratory, Kojinkai, Tsutsujigaoka, Miyagino-ku, Sendai, Japan

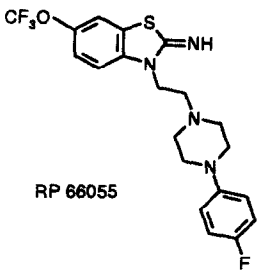
Abstract: The Synthesis of the enzymatically stable thymic humoral factor-γ2 analog is reported.

Ac-Leu-Glu-Asp-Gly-Pro-Lys-Phe-Leu-CH_2Cl

BioMed. Chem. **1994**, *2*, 793

SYNTHESIS, ANTICONVULSANT AND NEUROPROTECTIVE ACTIVITIES OF RP 66055, A RILUZOLE DERIVATIVE

P. Jimonet,* M. Barreau, J.-C. Blanchard, A. Doble, P. Laduron, C. Malgouris, O. Piot, J. Pratt, J. Rataud, M. Reibaud, S. Mignani and J.-M. Stutzmann
Rhône-Poulenc Rorer, Central Research, Centre de Recherche de Vitry-Alfortville, 94403 Vitry sur Seine Cedex, France



RP 66055

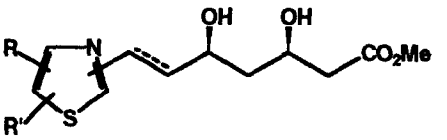
RP 66055, a riluzole derivative, has been characterized as a potent anticonvulsant and *in vivo* neuroprotective agent.

BioMed. Chem. **1994**, *2*, 799

7-(DISUBSTITUTEDTHIAZOLYL)-3,5-DIHYDROXY-6-HEPTENOIC-/HEPTANOIC ACID DERIVATIVES AS HMG-CoA REDUCTASE INHIBITORS

Violetta Cecchetti,[a] Arnaldo Fravolini,[a]* Pier Giuseppe Pagella,[b] Oriana Tabarrini,[a] and Andrea Temperini[a]
[a]*Istituto di Chimica Farmaceutica e Tecnica Farmaceutica, Università di Perugia, 06123 Perugia, Italy*
[b]*Mediolanum Farmaceutici, via S. G. Cottolengo 31, 20143 Milano, Italy*

A series of disubstituted thiazole, functionalized with the essential 3,5-dihydroxy-6-heptenoic or heptanoic chain, were prepared and evaluated in vitro as HMG-CoA reductase inhibitors. All the synthesized compounds showed a moderate inhibitory potency.



R and R' = *i*-Pr, C_6H_5, 4-F-C_6H_5

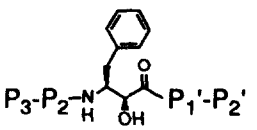
BioMed. Chem. **1994**, *2*, 807

STRUCTURE-ACTIVITY RELATIONSHIPS OF HIV-1 PR INHIBITORS CONTAINING AHPBA.

Mitsuya Sakurai,[a,*] Susumu Higashida,[a] Machiko Sugano,[a] Tomoaki Komai,[b] Ryuichi Yagi,[b] Yuji Ozawa,[b] Hiroshi Handa,[c] Takashi Nishigaki,[b] and Yuichiro Yabe,[a,*] [a]*Exploratory Chemistry Research and* [b]*Biological Research Laboratories, Sankyo Co. Ltd., 1-2-58 Hiromachi, Shinagawa-ku, Tokyo 140, and* [c]*Faculty of Bioscience and Biotechnology, Tokyo Institute of Technology, Nagatsuta, Midori-ku, Yokohama 227, Japan.*

Systematic replacement of the compounds containing AHPBA, at the P_3-P_2' sites, gave several potent and selective HIV-1 PR inhibitors.

AHPBA = 3-amino-2-hydroxy-4-phenylbutanoic acid

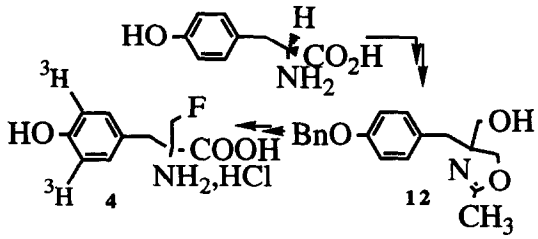


BioMed. Chem. **1994**, *2*, 827

Synthesis of [3',5'-$^3H_2$]-α-Fluoromethyl-Tyrosine as a Radioactive Specific Label of Rat Brain Tyrosine Hydroxylase

Pierre Lafargue[a], Alain Dodi[a], Michel Ponchant[a], Christine Garcia[b], Marion Le Cavorsin[b], Jean-François Pujol and Jean-Paul Lellouche*[a]
a) CE-Saclay, Bt 547, DBCM / SMM, 91191 Gif-Sur-Yvette, France.
b) CNRS-UCB UMR 105, rue G. Paradin, 69008 Lyon, France.

The [3',5'-$^3H_2$]-α-fluoromethyl-tyrosine **4** (specific activity 15.0 Ci/mmol) has been synthesized as a potentially useful radioactive probe for rat neuronal tyrosine hydroxylase.



BioMed. Chem. **1994**, *2*, 837

Cloning, Overexpression and Isolation of the Type II FDP Aldolase from *E. coli* for Specificity Study and Synthetic Application

L. Henderson, E. Garcia-Junceda, K. K.-C. Liu, Y.-L. Chen, G.-J. Shen, C.-H. Wong*
Department of Chemistry, The Scripps Research Institute, 10666N. Torrey Pines Road, La Jolla, CA 92037, USA.