

References

- Abbott, F.S., Kassam, J., Orr, J.M., and Farrell, K. The effect of aspirin on valproic acid metabolism. Clin pharmaco Ther. 40(1986): 94-100.
- Adkison, K.D.K., Ojemann, G.A., Report, R.L., and Shen, D.D. Distribution of unsaturated metabolites of valproate in human and rat brain-pharmacologic relevance?. Epilepsia. 36(8)(1995): 772-782.
- Alldredge, K.B. Seizure disorder. In T. Herfindal, D. Gourley and L.L. Heart (eds.), Clinical Pharmacy and Therapeutics, (5th ed.), U.S.A. : Willium & Wilkins, 1992.
- Annegers, J.F., Hauser, W.A., Shirts, S.B., and Kurland, L.T. Factors prognostic of unprovoked seizures after febrile convulsions. N Engl J Med. 316(1987): 413-498.
- Benveniste, H., and Huttemeier, P.C. Microdialysis; Theory and Application. Prog Neurobio. 35(1990): 195-215.
- Bialer, M., Hai-Yehia, A., Badir, K., and Hadad, S. Can we develop improved derivatives of valproic acid? Pharmacy World & Science. 16(1994): 2-5.
- Biggs, C.S., Peace, B.R., Whitton, P.S. The effect of sodium valproate on extracellular GABA and other amino acids in the rat ventral hippocampus : an in vivo microdialysis study. Brain Res. 594(1992): 138-142.
- Boonardt Saisorn, Chamnan Patarapanich, and Wichan Janwitayanuchit. Synthesis of monoureide analogues of valproic acid. Thai J. Pharm. Sci. 16(1992): 145-150.
- Bourgeois, B., et al. Felbamate:a double-blind controlled trial in patients undergoing presurgical evaluation of patial seizures. Neurology. 43(1993): 693-696.

- Bowery, N.G., Hill, D.R., and Moratalla, R. Neurochemistry and autoradiography of GABA_B receptor in mammalian brain : second-messenger system(s). In, E.A. Barnard and E. Costa (eds.), Allosteric modulation of amino acid receptors, pp. 59-172. New York: Raven Press, 1989.
- Browning, R.A. Overview of neurotransmission :Relationship to the action of antiepileptic drugs. In, C.L. Faingold and G.H. Fromn (eds.), Drug for control of epilepsy, pp. 23-56. Florida : CRC Press, 1991.
- Chapman, A., Keane P.E., Meldrum, B.S., Simiand, J., Vernieres, J.C. Mechanism of anticonvulsant action of valproate. Progress in Neurology. 19(1982): 315-359.
- Cooper, J.R., Bloom, F.E., and Roth, R.H. The biological basis of neuropharmacology, (6th ed.), pp. 133-189. New-York : Oxford University Press, 1991.
- Crawford, P.,et al. The lack of effect of sodium valproate on the pharmacokinetics of oral contraceptive steroids. Contraception. 33(1986): 23-29.
- Davidoff, R.A. Studies of neurotransmitter action (GABA, glycine and convulsants). In, A.A. Ward, J.K. Penny, and D.P. Purpura (eds.), Epilepsy : Association for research in nervous and mental disease, Vol 61, pp. 53-85. New York : Raven Press, 1983.
- Davies, J., and Richens, A. Neuropharmacology. In J. Laidlaw, A. Richens and D. Chadwick, Textbook of Epilepsy, (4th ed.), pp.1-19. London Churchill Livingstone, 1993.
- Davis, R., Peters, D.H., and McTavish, D. Valproic acid: A reappraisal of its pharmacological properties and clinical efficacy in Epilepsy. Drug. 47(2) (1994): 332-372.
- Diem, K. and Lentner, C. Scientific tables. (7 th ed.), pp. 54-55. Germany : Ciba Geigy Limited; 1972.

- Dreifuss, F. and Langer, D.H. Side effects of Valproate. Am. J. Med. 84(suppl 1A) (1988): 34-40.
- Dunham, N.W. and Miya, T.S. A note on a simple apparatus for detecting neurological deficit in rats and mice. J. Am. Pharm. Assoc. 46 (1957): 208-209.
- During, M.J., and Spencer, D.D. Extracellular hippocampal glutamate and spontaneous seizure in the conscious human brain. Lancet. 341(1993): 1607-1610.
- Eadie, M.J., Mckinnon, G.E., Dunstan, P.R., MacLaughlin, D., and Dickinson, R.G. Valproate metabolism during hepatotoxicity associated with the drug. Quart. J. Med. 284(1990): 1229-1240.
- Faden, A.L., Ellison, J.A., and Noble, L.J. Effect of competitive NMDA receptor antagonists in spinal cord injury. Eur. J. Pharmacol. 175(1990): 165-174.
- Farrent, M and Webster, R.A. Neuronal activity, amino acid concentration and amino acid release in the substantia nigra of the rat after sodium valproate, Brain Res. 504(1989) 49-56.
- Faught, E., et al. Felbamate monotherapy for partial-onset seizures : on active control trial. Neurology. 43(1993): 688-692.
- Fukuzako, H., and Izumi, K. Clinical aspects of the epilepsies. In G.Tunnicliff, and B.U. Raess (eds.), GABA mechanism in epilepsy, pp.1-30. New York: Wiley-Liss, 1991.
- Godin, Y., Heiner, L., Mark, J., and Mandle, P. Effects of di-n-propylacetate an anticonvulsive compound, on GABA metabolism. J. Neurochem. 16 (1969): 869-873.
- Goldring, S. Pathophysiology of epileptic discharge. In, S.G. Eleasson, A.L. Prensky, and W.B. Hardin, Jr. (eds.), Neurological pathophysiology, pp. 183-195. New York: Oxford University, 1978.

- Goulden, K.J., Dooley J.M., Camfield, P.R., and Fraser, A.D. Clinical valproate toxicity induced by acetylsalicylic acid Neurology. 37(1987): 1392-1394.
- Graves, N.M., Leppik, I.E. Antiepileptic medications in development. Ann Pharmacother. 25(1991): 978-986.
- Greenamyre, J.T., and Porter, R.H.P. Anatomy and Physiology of glutamate in the CNS. Neurology. 44(suppl 8)(1994): S₇-S₂₃.
- Hauser, W.A. The natural history of febrile seizures. In K.B., Ellenberg J.H. (eds.), Febrile seizures, pp . 5-17. New York: Reven Press, 1981.
- Hauser, W.A., Annegers, J.F., and Anderson, V.E. Epidemiology and the genetic epilepsy. In A.A. Ward, J.K. Penry, and D.P. Purpura (eds.), Epilepsy Research Publications: Association for Research in Nervous and Mental Disease Volume 61, pp. 225-237. New York: Reven Press, 1983.
- Hopkin, A. Clinical Neurology, pp. 186-207. New York: Oxford University Press, 1993.
- Horton, R.W. GABA dysfunction in animal models of epilepsy. In G. Tunnicliff, and B.U. Raess, (eds.), GABA mechanisms in epilepsy, pp. 121-147. New York: Wiley-Liss, 1991.
- Ichinose, N., Nakamura, K., Shimizu, C., Kurokura, H., and Okamoto, K. High-performance liquid chromatography of 5,8,11,14,17-eicosapentanoic acid in fatty acid (C₁₈ and C₂₀) by labeling with 9-anthyl diazomethane as a fluorescent agent. J. Chromatogr. 295(1984): 463-469.
- Jeavons, P.M. Valproate: Toxicity, In D.M. Woodbury, J.K. Penry, and C.E. Pippenger (eds.), Antiepileptic drugs, 2nd ed, pp. 601-610. New York: Raven Press, 1982.
- Johnson, T. Valproate versus carbamazepine for seizures. N Engl J Med. 328(1993): 208.

- Ladarola, M.J. and Gale, K. Cellular compartments of GABA in brain and their relationship to anticonvulsant activity. Mol. Cell Biochem. 39(1981):305-330.
- Lamb, R.J., Miller, A.A. Effect of lamotrigine and some known anticonvulsant drugs on visually-evoked after discharge in the conscious rat. Br J Pharmacol. 86(1985): 765.
- Leach, M.J., Marden, C.M., and Miller A.A. Pharmacological studies on lamotrigine, a novel potential antiepileptic drug:II Neurochemical studies on the mechanism of action. Epilepsia. 27(1986): 490-497.
- Levitin, E.S., et al. Structural and functional basis for GABA_A receptor heterogeneity. Nature. 335(1988): 76-79.
- Levy, R.H., and Koch, K.M. Drug interactions with valproic acid. Drug. 24(1982): 543-556.
- Lindroth, P., and Mopper, K. High performance liquid chromatographic determination of subpicromole derivatization of amino acids by precolumn fluorescence derivatization o-phthalaldehyde. Anal. Chem. 51(1979): 1667-1674.
- Litchfield, J. T., and Wilcoxon, F. W. A simplified method of evaluating dose-effect experiments. J Pharmacol. Exp. Ther. 96(1949): 99-119.
- Loscher, W. Valproate induced change in GABA metabolism at the subcellular level. Biochem Pharmacol. 30(1981): 1364-1366.
- and Bohme,G., Schafer, H., and Kochen, W. Effect of metabolites of valproic acid on the metabolism of GABA in brain and brain nerve endings. Neuropharmacology. 20(1981): 1187-1192.
- and Nolting, B. The role of technical, biological and pharmacological factors in the laboratory evaluation anticonvulsant drugs. IV. Protective indices Epilepsy Res. 9(1991): 1-10.

- .and Vetter, In vivo effects of aminoxyacetic acid and valproic acid on nerve terminal (synaptosomal) GABA levels in discrete brain regions of the rat brain: correlations to pharmacological activities, *Biochem Pharmacol.* 34 (1985): 1747-1756.
- .Nolting, B., and Fassbender, C. The role if technical, biological and pharmacological factors in the laboratory evaluation of anticonvulsant drugs. I. The influence of administration vehicles. *Epilepsy Res.* 7(1990): 173-181.
- Matsumoto, R.R. GABA receptors are cellular differences reflected in function. *Brain Research Reviews.* 14(1989): 203-225.
- Matsuo, F., et al. Placebo-controlled study of the efficacy and saftety of lamotrigine in patients with partial seizures. *Neurology.* 43(1993): 2284-2291.
- Mattson, R.H.,et al. A comparison of valproate with carbamazepine for the treatment of complex partial seizures and secondary generalized tonic-clonic seizures in adults. *N Engl J Med.* 327(1992): 765-771.
- McGeer, P.L., Eccles, J.C., and McGeer, E.G. Molecular neurobiology of the brain, pp. 151-173. New York: Plenum Press, 1987.
- .Eccle, J.C., and McGeer, E.G. Molecular neurobiology of the mammalian brain, (2 nd. ed.), pp. 175-234. New York: Plenum Press, 1988.
- Meldrum, B.S. Excitatory amino acid transmitter in epilepsy. *Epilepsia.* 35 (suppl 2) (1991):S₁-S₃.
- .The role of glutamate in epilepsy and other CNS disorders. *Neurology.* 44 (suppl 8)(1994): S₁₄-S₂₃.
- .Chapman,A.G., Patel,S., Swan, J. Competitive NMDA antagonists as drugs. In C.J., Watkins and L.G., Collingridge (eds.), The NMDA receptor, pp. 207-216. Oxford: Oxford University Press, 1989.

- Menkes, J.H. Textbook of child neurology, 4th ed., pp.602-674. Philadelphia: Lea & Fibiger, 1990.
- Messenheimer, J., et al. Lamotrigine therapy for partial seizure : a multicenter, placebo controlled, double-blind, cross-over trial. Epilepsia. 35(1994): 113-121.
- Millan, M.H., Chapman, A.G., and Meldrum, B.S. Extracellular amino acid level in hippocampus during pilocarpine-induced seizures. Epilepsy Res. 14(1993): 139-148.
- Moore, M., Keane, P.E., Verniteres, J.C., Simiand, J., Roncucci, R. Valproate: recent findings and perspectives. Epilepsia. 25(suppl 1)(1984): S5-S9.
- Oller, L.F., Russi, A., Daurella, L.O. Lamotrigine therapy in Lennox-Gastaut syndrome. Epilepsia. 32(suppl 1)(1991): 58.
- Olsen, R.W., and Tobin, A.J. Moleccula biology of GABA_A receptors. FASEB Journal. 4(1990): 1469-1480.
- Orr, J.M., et al., Interaction between valproic acid and aspirin in epileptic chlidren; serum protein binding and metabolic effects. Clin Pharmacol Ther. 31(1982) : 642-649.
- Pellegrino, J.L., Pellegrino, A.S., and Crushman, A.J. A stereotaxic atlas of the Rat Brain. New York: Plenum Pressation, 1979.
- Perucca, E., et al. Pharmacokinetics of valproic acid in the elderly. BR J CLIN PHARMACO. 17(1984): 665-669.
- Phillips, N.I., and Fowler, L.J. The effect of sodium valproate on γ -amino butyrate metabolism and behavior in native and ethanolamine-o-sulfate pretreated rats and mice. Biochem Pharmacol. 3(1982): 2257-2261.
- Pitkanen, A., Saano, V., Hyvonen,k., Airaksinen, M.M., and Riekkinen, P.J. Decressed GABA, benzodiazepine, and picrotoxin receptor binding in brains of rat after cobalt-induced epilepsy. Epilepsia. 28(1987): 11-16.

- Ponchulee Supatchipisit. Anticonvulsant activity and effects of (N-hydroxymethyl)-2-propylpentamide on the central nervous system. Master's Thesis, Chulalongkron University, 1995.
- Porter, R.J. Classification of epileptic seizures and epileptic syndromes. In, J. Laidlaw, A. Richens, and D. Chadwick, Textbook of epilepsy, (4th ed.), pp 1-19 London: Churchill Livingstone, 1993.
- Rall, T.W., and Schleifer, L.S. Drugs effective in the therapy of the epilepsies. In, A.G. Gilman (eds.), Goodman and Gilman's the Pharmacological basic of therapeutics, (5th ed.), pp. 436-462. New York: Pergamon Press, 1990.
- Ribak, C.E. Morphological, biochemical, and immunocytochemical changes of cortical, GABAergic system in epileptic foci. In, A.A. Ward, J.K. Penry, and D.P. Pupura (eds.), Epilepsy Research Publications: association for research in nervous and mental disease. Volume 61, pp. 109-130. New York: Raven Press, 1983.
- Ribak, C.E., Harris, A.B., Vaughn, J.E., and Roberts, E. Inhibitory GABAergic nerve terminals decrease at sites of focal epilepsy. Science 205(1979): 211-214.
- Rogawski, M.A., and Porter, R.J. Antiepileptic drugs: Pharmacological mechanism and clinical efficacy with consideration of promising developmental stage compounds. Pharmacol. Rev. (1990): 223-286.
- Sachdeo, R., et al. Felbamate monotherapy: controlled trial in patients with partial onset seizures. ANN NEUROL. 32(1992): 386-392.
- Schwartzkroin, P.A. Basic mechanisms of epileptogenesis. In, E.Wyllie (eds.), The treatment of epilepsy: principle and practice, pp. 83-98. Philadelphia: Lea & Fibiger, 1993.

- Silverman, R.B., Andruszkiewicz,R., Nanavati, S.M., Taylor, C.P., and Vartanian, M.G. 3-akyl-4-aminobutyric acids: the first class of anticonvulsant agents that activates L-glutamic acid decarboxylase, *J.Med.Chem.* 34(1991) 2295-2298.
- Sivenius,J., et al. Double-blind study of gabapentin in the treatment of partial seizures. *Epilepsia*. 32(1991): 539-542.
- Slater, G.E., and Johnston, G.D. Sodium valproate increases potassium conductance in Aplysia neurons. *Epilepsia*. 19(1978): 379-384.
- Stagnitto, M.L., et al. Preclinical profile of macemide: a novel anticonvulsant effective against maximal electroshock seizures in mice *Epilepsy Res.* 7 (1990): 11-28
- Steiner, T.J., Yuen, A.W. Comparison of lamotrigine and phynetoin monotherapy in newly diagnosed epilepsy. *Epilepsia*. 35(suppl 8) (1994) :31.
- Sugimoto, T., Woo, M., Nishida, N., Takeuchi, T., Sakane, Y., and Kobayashi, Y. Hepatotoxicity in rat following administration of valproic acid. *Epilepsia* 28 (1987): 142-146.
- Swinyard, E.A., and Woodhead, J.H. Experimental detection, quantification and evaluation of anticonvulsants. In, D.M. Woodbury, J.K. Penry and C.E. Pippenger (eds.), *Antiepileptic Drug*, pp. 111-126. New York: Revan Press, 1982.
- Thongchai Sooksawate. Anticonvulsant effects of N-(2-propylpentanoyl)Urea. Master's Thesis, Chulalongkron University, 1995.
- Toman, J.E.P., Swinyard, E.A. and Goodman, L.S. Properties of maximal seizures and their alteration by anticonvulsant drugs and other agents. *J.Neurophy.* 9(1946): 231-240.

- Van der Laan, J.W., De boer T. and Bruinvels. Di-n-propylacetate and GABA degradation, Preferential inhibition of succinic seminase. J Neurochem. 32(1979):1769-1780.
- Vanyer, P., Cash, C.D. and Maitre, M. Is the anticonvulsant mechanism of valproate linked to its interaction with the cerebral γ -hydroxy butyrate system? Trends in Pharmacology and Therapeutics. (1988): 127-129.
- Whittle, Sr. and Turner, S.J. Effects of the anticonvulsant sodium valproate on γ -aminobutyrate and aldehyde metabolism in brain. J Neurochem. 31(1978): 1453-1459.
- Wilder, B.J. The treatment of epilepsy. Neurology. 45(Suppl 2)(1995); S₇.
- Wojcik, W.J., Paez, X., and Ulivi, M. A transduction mechanism for GABA_B receptors. In E.A. Barnard and E.Costa (eds.), Allosteric modulation of amino acid receptors, pp. 173-193. New York: Reven Press, 1989.
- Yuen, A.W. Lamotrigine VS carbamazepine as monotherapy in patient with newly diagnosed or recurrent epilepsy. Epilepsia 35 (suppl 8)(1994): 31.
- Zeise, M.L., Lasparow, S., and Zieglgansberger W. Valproate suppresses N-methyl-D-aspartate evoked, transient depolarizations in the rat neocortex in vitro. Brain res. 554 (1991): 345-348.
- Zucker, R.S., and Lando, L. Mechanism of transmitter release: Voltage hypothesis and calcium hypothesis. Science. 231(1986): 574-579.

Appendices

Total amount of aspartate within 180 min after injection

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	36	90.40	18.48	3.08
PEG400	47	99.60	48.74	7.11
VPA200	44	91.07	27.19	4.10
VPA400	50	88.55	57.77	8.17
VPU200	45	85.63	28.24	4.21
VPU400	44	93.63	34.29	5.17

Total amount of glutamate within 180 min after injection

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	36	139.38	17.4	2.90
PEG400	50	125.87	42.14	5.96
VPA200 ^{a,b,d,f}	43	86.33	40.72	6.21
VPA400	43	141.02	59.67	9.10
VPU200 ^{a,b,d,f}	53	80.91	25.04	3.44
VPU400 ^{a,d}	44	116.07	64.01	9.65

^a P<0.05 denotes statistically significant from NSS

^b P<0.05 denotes statistically significant from PEG400

^d P<0.05 denotes statistically significant from VPA400

^f P<0.05 denotes statistically significant from VPU400

Total amount of glycine within 180 min after injection

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	36	86.16	17.4	2.90
PEG400	50	65.78	42.14	5.96
VPA200 ^a	45	65.47	61.04	9.10
VPA400 ^{a,b,c,e,f}	51	161.88	44.34	6.21
VPU200	54	80.55	70.91	9.65
VPU400	36	84.24	20.64	3.44

^a P<0.05 denotes statistically significant from NSS

^b P<0.05 denotes statistically significant from PEG400

^c P<0.05 denotes statistically significant from VPA200

^d P<0.05 denotes statistically significant from VPU200

^e P<0.05 denotes statistically significant from VPU400

Total amount of GABA within 180 min after injection

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	36	86.78	37.2	6.2
PEG400	50	81.27	33.65	4.76
VPA200 ^{a,b,e}	45	143.70	97.60	14.55
VPA400 ^{a,b,e,f}	53	169.70	187.68	25.78
VPU200 ^{c,d}	53	94.16	63.26	8.69
VPU400 ^c	45	107.61	72.98	10.88

^a P<0.05 denotes statistically significant from NSS

^b P<0.05 denotes statistically significant from PEG400

^c P<0.05 denotes statistically significant from VPA200

^d P<0.05 denotes statistically significant from VPA400

^e P<0.05 denotes statistically significant from VPU200

^f P<0.05 denotes statistically significant from VPU400

Amount of aspartate at various time

Time 1 (20 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	89.9	26.64	13.32
PEG400	6	79.23	47.74	19.49
VPA200	5	98.50	17.06	7.63
VPA400	6	89.96	57.93	23.65
VPU200	5	90.34	30.65	13.71
VPU400	5	80.11	34.74	15.54

Time 2 (40 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	99.1	51.18	25.59
PEG400	6	87.86	51.70	21.11
VPA200	5	104.96	27.54	12.32
VPA400	5	121.49	119.18	53.30
VPU200	5	71.97	30.47	13.63
VPU400	5	103.07	55.09	24.64

Time 3 (60 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	82.01	27.46	13.73
PEG400	5	79.39	38.88	17.39
VPA200	5	93.35	27.97	12.51
VPA400	6	130.08	121.56	49.63
VPU200	5	78.70	38.52	17.23
VPU400	5	100.93	56.50	25.27

Time 4 (80 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	107.02	52.39	26.19
PEG400	5	101.08	51.54	23.05
VPA200	4	69.62	17.42	8.71
VPA400	6	84.11	74.80	30.54
VPU200	5	83.08	25.08	11.22
VPU400	5	90.18	68.44	30.61

Time 5 (100 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	93.54	31.64	15.82
PEG400	5	126.24	84.24	37.66
VPA200	5	82.29	18.96	8.48
VPA400	6	92.43	84.40	34.46
VPU200	5	79.63	31.72	14.19
VPU400	5	78.56	31.64	14.15

Time 6 (120 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	93.15	72.16	36.08
PEG400	5	123.08	63.48	28.39
VPA200	5	80.50	17.82	7.97
VPA400	5	85.16	40.24	18.00
VPU200	5	106.22	65.89	29.47
VPU400	5	76.25	33.70	16.85

Time 7 (140 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	87.81	46.78	23.39
PEG400	5	76.13	35.84	16.03
VPA200	5	85.75	23.85	10.67
VPA400	6	65.61	20.20	8.25
VPU200	5	67.79	22.71	10.16
VPU400	5	87.33	42.41	18.97

Time 8 (160 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	91.18	60.04	30.02
PEG400	5	98.87	44.11	19.73
VPA200	5	108.03	52.90	23.66
VPA400	5	58.76	11.98	5.36
VPU200	5	97.00	67.12	30.02
VPU400	5	125.31	40.80	18.25

Time 9 (180 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	74.96	32.58	16.29
PEG400	5	129.24	57.82	25.86
VPA200	5	91.24	49.46	22.12
VPA400	5	65.30	20.92	9.32
VPU200	5	95.93	93.51	41.82
VPU400	5	98.11	20.16	9.02

Amount of glutamate at various time

Time 1 (20 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	143.46	49	24.50
PEG400	6	120.57	56.68	23.14
VPA200	5	86.21	27.43	12.27
VPA400	5	112.77	57.51	25.72
VPU200	6	97.08	55.84	22.80
VPU400	5	92.84	17.97	8.04

Time 2 (40 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	121.36	24.46	12.23
PEG400	6	136.40	30.27	12.36
VPA200	5	89.87	39.71	17.76
VPA400	5	132.95	61.67	27.58
VPU200	6	92.70	39.90	16.29
VPU400	5	96.03	58.09	25.98

Time 3 (60 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	151.33	77.36	38.68
PEG400	6	143.91	44.21	18.05
VPA200	4	66.78	11.04	5.52
VPA400	5	165.98	147.98	66.18
VPU200	6	116.39	59.25	24.19
VPU400	5	109.12	66.34	29.67

Time 4 (80 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	108.70	25.04	12.52
PEG400	6	112.87	276.47	11.44
VPA200	4	59.90	23.26	11.63
VPA400	5	153.73	99.14	44.34
VPU200	6	86.14	45.29	18.49
VPU400	5	105.21	63.28	28.30

Time 5 (100 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	139.21	66.78	33.39
PEG400	5	111.58	31.55	14.11
VPA200	5	77.69	38.72	17.32
VPA400	5	112.69	63.77	28.52
VPU200	6	74.26	40.00	16.33
VPU400	5	93.01	25.73	11.51

Time 6 (120 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	118.22	20.36	10.18
PEG400	6	147.32	107.16	43.75
VPA200	5	82.97	70.55	31.38
VPA400	4	174.41	117.16	58.58
VPU200	6	57.37	35.12	14.34
VPU400	4	111.37	40.52	20.26

Time 7 (140 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	147.01	56.12	28.06
PEG400	5	83.98	55.27	24.72
VPA200	5	106.33	66.47	29.73
VPA400	5	129.23	109.63	49.03
VPU200	6	57.23	20.60	8.41
VPU400	5	126.54	78.88	35.28

Time 8 (160 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	149.37	43.62	21.81
PEG400	5	138.16	46.26	20.69
VPA200	5	119.59	65.40	29.25
VPA400	4	150.7	174.68	87.34
VPU200	5	97.86	40.71	18.21
VPU400	5	158.56	146.90	65.70

Time 9 (180 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	175.82	71.30	35.65
PEG400	5	138.07	103.99	46.51
VPA200	5	87.65	35.26	15.77
VPA400	5	136.60	138.27	61.84
VPU200	6	79.97	21.67	8.85
VPU400	5	152.38	74.46	33.30

Amount of glycine at various time

Time 1 (20 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	96.26	36.84	18.42
PEG400	6	96.50	54.67	22.32
VPA200	5	90.62	60.44	27.03
VPA400	6	117.27	21.80	8.90
VPU200	6	88.27	19.62	8.01
VPU400	4	119.58	102.78	51.39

Time 2 (40 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	77.69	30.86	15.43
PEG400	6	73.39	37.13	15.16
VPA200	5	86.34	37.92	16.96
VPA400	5	149.42	69.04	30.88
VPU200	6	66.28	23.61	8.64
VPU400	4	69.91	40.28	20.14

Time 3 (60 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	79.07	38.78	19.39
PEG400	6	73.27	39.06	15.95
VPA200	5	47.92	20.21	9.04
VPA400	6	186.18	155.93	63.66
VPU200	6	87.09	45.90	18.74
VPU400	4	136.49	96.12	48.06

Time 4 (80 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	90.52	181.04	27.14
PEG400	6	93.69	77.30	31.56
VPA200	5	88.72	122.13	54.62
VPA400	6	137.17	102.11	41.69
VPU200	6	91.28	22.14	9.04
VPU400	4	63.44	26.06	13.03

Time 5 (100 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	79.66	42.56	21.28
PEG400	5	60.83	33.78	15.11
VPA200	5	60.68	38.84	17.37
VPA400	6	173.10	146.62	59.86
VPU200	6	71.93	15.70	6.41
VPU400	4	72.30	29.90	14.95

Time 6 (120 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	88.09	66.34	33.17
PEG400	6	62.02	57.19	23.35
VPA200	5	43.91	24.64	11.02
VPA400	5	143.54	72.33	32.35
VPU200	6	72.74	20.50	8.37
VPU400	4	76.34	4.74	2.37

Time 7 (140 min after injection)

Group	count	Mean	Standard Diviation	Standard Error of Mean
NSS	4	84.10	53.48	26.74
PEG400	5	36.42	17.37	7.77
VPA200	5	55.22	29.16	13.04
VPA400	6	164.56	110.32	45.04
VPU200	6	69.60	26.06	10.64
VPU400	4	66.67	34.00	17.00

Time 8 (160 min after injection)

Group	count	Mean	Standard Diviation	Standard Error of Mean
NSS	4	78.11	51.38	25.69
PEG400	5	60.57	37.63	16.83
VPA200	5	66.85	10.21	4.57
VPA400	5	182.20	150.80	67.44
VPU200	6	84.24	36.81	15.03
VPU400	4	72.88	33.64	16.82

Time 9 (180 min after injection)

Group	count	Mean	Standard Diviation	Standard Error of Mean
NSS	4	101.94	100.66	50.33
PEG400	5	63.26	54.29	24.28
VPA200	5	48.98	23.43	10.48
VPA400	6	203.52	72.48	29.59
VPU200	6	92.98	62.21	25.40
VPU400	4	103.81	47.82	23.91

Amount of GABA at various time

Time 1 (20 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	108.60	37.20	18.60
PEG400	6	92.48	43.38	17.71
VPA200	5	155.70	82.51	36.90
VPA400	6	87.46	28.73	11.73
VPU200	6	90.85	38.33	15.65
VPU400	5	87.47	83.56	37.37

Time 2 (40 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	74.96	20.04	10.02
PEG400	6	80.37	36.96	15.09
VPA200	5	98.44	55.76	24.94
VPA400	6	120.59	80.04	32.68
VPU200	6	79.30	30.17	12.32
VPU400	5	71.95	47.38	21.19

Time 3 (60 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	56.27	17.52	8.76
PEG400	6	85.58	50.09	20.45
VPA200	5	92.57	44.34	19.83
VPA400	6	91.21	70.25	28.68
VPU200	6	90.89	43.06	17.58
VPU400	5	139.48	96.46	43.14

Time 4 (80 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	68.89	13.46	6.73
PEG400	6	68.24	48.91	19.97
VPA200	5	105.62	90.82	40.62
VPA400	6	136.67	129.33	52.84
VPU200	6	81.32	30.69	12.53
VPU400	5	71.50	272.57	121.90

Time 5 (100 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	77.61	14.22	7.11
PEG400	5	64.20	23.03	10.30
VPA200	5	117.47	67.79	30.32
VPA400	6	128.52	114.24	46.64
VPU200	6	84.70	38.04	15.53
VPU400	5	137.18	150.15	67.15

Time 6 (120 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	92.69	51.58	25.79
PEG400	6	106.20	48.94	19.98
VPA200	5	148.10	104.44	46.71
VPA400	6	163.92	183.85	75.06
VPU200	6	89.47	56.77	23.18
VPU400	5	68.64	94.18	42.12

Time 7 (140 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	99.03	51.2	25.60
PEG400	5	63.36	25.8	11.54
VPA200	5	170.69	172.17	77.90
VPA400	6	272.83	272.94	111.43
VPU200	6	65.64	24.88	10.16
VPU400	5	68.43	61.22	27.38

Time 8 (180 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	91.47	49.14	24.57
PEG400	5	90.61	45.10	20.17
VPA200	5	184.10	201.84	90.27
VPA400	6	236.00	294.69	120.30
VPU200	5	156.66	72.58	32.46
VPU400	5	135.37	151.00	67.53

Time 9 (180 min after injection)

Group	count	Mean	Standard Deviation	Standard Error of Mean
NSS	4	111.66	52.06	26.03
PEG400	5	80.46	64.33	28.77
VPA200	5	220.67	282.86	126.50
VPA400	5	290.89	334.82	149.74
VPU200	6	108.07	91.02	37.16
VPU400	5	138.07	145.54	65.09

Curriculum vitae

Miss Tipsuchon chunngam was born on 19 th October, 1971, in Lopburi, Thailand. She had graduated with Bachelor Degree in Pharmacy in 1994 from faculty of Pharmaceutical Sciences, Rangsit University, Phatomthani, Thailand. After graduation, she continue studies the master's Degree programme in Pharmaceutical Sciences at Chulalongkorn University.

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