

REFERÊNCIAS BIBLIOGRÁFICAS

Agência Nacional de Vigilância Sanitária (Brasil). Resolução RDC nº 136, de 29 de maio de 2003. Dispõe sobre o registro de medicamento novo. *Diário Oficial da República Federativa do Brasil*, Brasília, 02 jun 2003.

AHEER F., EL- NAHAS F.A, EL ASHWY IM. Reproductive and Cytogenetic Toxicity of Metronidazole in Male Mice. *Basic Clinical Pharmacology Toxicology*, Egypt, 94: 226-231, 2004.

ANGOSTO, M. C.; GÓMES-LECHÓN, M. J. *Citocromo P450*. 1. ed. Madri: Real Academia Nacional de Farmácia, 2004. 494 p.

AL-HABORI M, AL-AGHBARI A, AL-MAMARY M, BAKER M . Toxicological evaluation of *Catha edulis* leaves: A long term feeding experiment in animals. *Journal of Ethnopharmacology*, Ireland, 83: 209-217,. 2002

ALVAREZ-GUERRA M, HANNAERT P, HIDER H, CHIAVAROLI C, GARAY RP. Vascular permeabilization by intravenous arachidonate in the rat peritoneal cavity: antagonism by antioxidants. *European Journal of Phamacology*, Netherlands, 466:199-205, 2003

AMACHER E.D.; ADLER R.; HERATH A.; TOWNSEND R.R. Use of proteomic methods to identify serum biomarkers associated with rat liver toxicity or hypertrophy. *Clinical Chemistry*, 51: 10 1796-1803, 2005.

BAHR V., ULLMANN U., 1983. The influence of metronidazole and its two metabolites on murine *in vitro* lymphocyte transformation. *Eur. J. Clin. Microbiol* 2: 568-570.

BENDESKY A, MENÉNDEZ D., WEGMAN P.. Is metronidazole carcinogenic? *Mutation Research*, Mexico, 511: 133-144,, 2002

BOTHAM, P. A. Acute systemic toxicity – prospects for tiered testing strategies. *Toxicology in Vitro*, England, v. 18, n. 2, p. 227-230, 2004.

BRITO, A. S. *Manual de Ensaios Toxicológicos in vivo*. 1. ed. Campinas: UNICAMP, 1994. 122 p

BUSATTI H. *Investigação do potencial giardicida de análogos do metronidazol.* Instituto de Ciências Biológicas, Universidade Federal de Minas Gerais, Belo Horizonte, 32 pp. 2006

CAVALCANTI JCM, ABREU FC, OLIVEIRANV, MOURA MBF, CHAVESJG, ALVESRJ, BERTINARIA M, FRUTTERO R, GOULART MF. Effect of the Leaving Group on the Eletrodic Reduction Mechanism of Anti- Helicobacter pylori Metronidazole derivates, in Aprotic and Protic Media. *Bioelectrochemistry*, Maceió, 63:353-357, 2004

DESCOTES, J. From clinical to human toxicology: linking animal reserch and risck assesment in man. *Toxicology Letters*, Netherlands, v. 149, p.3-10, 2003.

DUNN, B.E.; COHEN H.; BLASER, M. J. *Helicobacter pylori.* *Clinical Microbiology.* 10:720-41,. 1997

EATON, D. L.; KLAASSEN, C. D. Principles of toxicology. In: KLAASSEN, C. D. *Casarett and Doull's Toxicology: The Basic Science of Poisons.* 5. ed. New York: McGraw-Hill, 1996. Chapter 2, p. 13-33.

ELIZONDO G, MONTERO R, HERRERA JE, HONG E, OSTROSKY-WEGMAN P.. Lymphocyte proliferation kinetic and sister- chromatid exchanges in individuals treated with metronidazole. *Mutatation Research*, Mexico, 2: 133-137, 1994

ELIZONDO G., GONSEBATT, SALAZAR A.M.; LARES J.; SANTIAGO P.; HERRERA J.F.; HONG F.; OSTROSKY-WEGMAN P. 1996. Genotoxic effects of metronidazole. *Mutat. Res.* 370: 75-80.

ELLEDGE S.J.1996. Cell cycle checkpoints: preventing an identity crisis. *Science* 74: 1664-1672.

EEL-NAHAS FA, EL ASHMWY IM. Reproductive and Cytogenetic Toxicity of Metronidazole in Male Mice. *Basic Clinical Pharmacology Toxicology*, Egypt, 94: 226-231, 2004

FÉRES C.A.O.; *et al.* Acute and Chronic toxicological studies of *Dimorphandra mollis* in experimental animals. *Etinopharmacology*, Belo Horizonte. 108: 450-456, 2006.

FOOD AND DRUG ADMINISTRATION. CENTER FOR DRUG EVALUATION AND RESEARCH (CDER). Guidance for industry. Disponível em: <<http://www.fda.gov/cder/guidance/index.htm>> Acesso em: 09 de dez de 2006.

FOOD AND DRUG ADMINISTRATION (Food and Drug Administration). Center for Drug Evaluation and Research (CDER). Guidance for industry: Nonclinical Studies for Development of Pharmaceutical Excipients. September. 2002. Disponível em: <<http://www.fda.gov/cder/guidance/index.htm>> Acesso em: 13 de setembro 2005.

FRENCK, R.W. & CLEMENS, J. Helicobacter in the Developing World. *Microbes and Infection*, Cairo, 5, pg 705 – 713, 2003

FUJII, T. Toxicological correlation between changes in blood biochemical parameters and liver histopathological findings. *Journal of Toxicological Sciences*, United States, v. 22, n. 3, p. 161-183, 1997

GÁMEZ R, MAS R, NOA M, MENÉNDEZ R, ALEMÁN C, ACOSTA P, GARCIA H, HERNÁNDEZ C, AMOR A, PÉREZ J, GOICOCHEA E. Acute and oral subchronic toxicity of D-003 in rats. *Toxicology Letters*, Netherlands, 118: 31-41, 2000.

GRIFFIN G., STOKES W.S., PAKES S.P., GAUTHIER C. The ICLAS/CCAC International Symposium on Regulatory Testing and Animal Welfare. *ILAR Journal*, Ottawa, v. 43: S1-S4, 2002.

HARTWELL L.H., KASTAN M.B. 1994. Cell cycle control and cancer. *Science* 266: 1821-1828.

HASUMURA M., YASUHARA K., TAMURA T., IMAI T., MITSUMORI K., HIROSE M., 2004. Evaluation of the toxicity of enzymatically decomposed rutin with 13- weeks dietary administration to Wistar rats. *Food Chem Toxicol* 42: 439-444.

IVERSEN, P. O.; NICOLAYSEN, G. Water-for life. *Tidsskrift for den Norske Laegeforening*, Norway, v. 123, n. 23, p. 3402-3405, 2003.

JANBAZ, K. H.; SAEED, S. A.; GILANI, A. H. Protective effect of rutin on paracetamol- and CCl₄-induced hepatotoxicity in rodents. *Fitoterapia*, Netherlands, v. 73, n. 7-8, p. 557-563, 2002

JENKINS D.J.A. *et al.* Effects of antibiotics as cholesterol-lowering agents. *Metabolism Clinical and Experimental*, Tronto. 54: 103-112, 2005.

JENKINS KJ, HIDIROGLOU M. Effects of Elevated Iodine in Milk replacer on Calf Performance. *Journal of Dairy Science*, Ottawa 73: 804-807, 1990

JENKS, P.J. & EDWARDS, D.I. Metronidazole Resistance in *Helicobacter pylori*. *International Journal of Antimicrobial Agents*, Nottingham, 19, p. 1-7, 2002.

KANEKO, J.J.; BRUSS, M.L. Clinical biochemistry of domestic animals, 5ed., San Diego: Academic Press, p. 932, 1997.

LADICS, G. S.; SMITH, C.; HEAPS, K.; ELLIOT, G. S.; SLONE, T. W.; LOVELESS, S. E. Possible incorporation of an immunotoxicological functional assay for assessing humoral immunity for hazard identification purposes in rats on standard toxicology study. *Toxicology*, Ireland, v. 96, n. 3, p. 225-238, 1995.

LAMP K.C., FREEMAN C., KLUTMAN N., LACY M.. Pharmacokinetic and pharmacodynamics of the Nitroimidazole Antimicrobials. *Clinical Pharmacokinetic*, United States, 36: 353-373, 1999.

MENÉNDEZ D *et al.* Role of P53 functionality in the genotoxicity of metronidazole and its hydroxy metabolite. *Mutation Research*, 501: 57-67, 2002.

MCCLAIN RM, DOWNING CJ, EDGCOMB EJ. Effect of Metronidazole on Fertility and Testicular Function in Male rats. *Fundamental Applied Toxicology*, New Jersey, 12: 386-396, 1989

MURALIDHARA, NARASIMHAMURTHY K, VISWANATHA S, RAMESH BS. Acute and subchronic toxicity assessment of Debitterized Fenugreek powder in the mouse and rat. *Food Chemical Toxicology*, England, 37: 831-838, 1999.

NAKAMURA H, FURUKAWA F, NISHIKAWA A, MIYAUCHI M, SON HY, IMAZAWA T. Oral toxicity of a tocotrienol preparation in rats. *Food Chemical Toxicology*, England, 39: 799-805, 2001

OECD. Guidelines for the testing of chemicals, n. 401: Acute Oral Toxicity. Paris, may, 1981.

OECD. Guidelines for the testing of chemicals, n. 407: *Repeated dose 28-day toxicity studies in rodents*. Paris, jul, 1995.

OECD. Guidelines for the testing of chemicals, n. 420: *Acute oral toxicity – fixed dose procedure*. Paris, dec, 2001.

OGA, S. *Fundamentos de Toxicologia*. 2. ed. São Paulo: Atheneu, 2003, 474 p.

OLSON, H.; BETTON, G.; ROBINSON, D.; THOMAS, K.; MONRO, A.; KOLAJA, G.; LILLY, P.; SANDERS, J.; SIPES, G.; BRACKEN, W.; DORATO, M.; DEUN, K. V.; SMITH, P.; BERGER, B.; HELLER, A. Concordance of Toxicity of Pharmaceuticals in Humans and in Animals. *Regulatory Toxicology and Pharmacology*, United States, v. 32, n. 1, p. 56-67, 2000.

PORTAL-CELHAY C, PEREZ-PEREZ GI. Immune responses to *Helicobacter pylori* colonization: mechanisms and clinical outcomes. *Clinical Science*, London 3: 305-314, 2006.

RAETHER W., HÄNEL H. Nitroheterocyclic drugs with broad spectrum activity. *Parasitology Research*, Frankfurt 90:S19-S39, 2003.

RAZA, M.; AL-SHABANAH, O. A.; EL-HADIYAH, T. M.; AL-MAJED, A. A. Effect of prolonged vigabatrin treatment on hematological and biochemical parameters in plasma, liver and kidney of Swiss albino mice. *Scientia Pharmaceutica*, Austria, v. 70, n. 2 , p. 135-145, 2002.

SAMPAIO, I. B. M. *Estatística Aplicada à Experimentação Animal*. 2. ed. Belo Horizonte: Fundação de Ensino e Pesquisa em Medicina Veterinária e Zootecnia, 2002. 265 p.

SANYAL S. N., JAMBA L., CHANNAN M. Effect of the Antiprotozoal Agent Metronidazole (Flagyl) on Absorptive and Digestive Functions of the Rat Intestine. *Annales of Nutrition and Metabolism*, India, 36: 235-243, 1992.

SHAMKHANI K., AZARPIRA M., AKBAR M.H.. An open label crossover trial of effects of metronidazol on hyperlipidaemia. *International Journal of Cardiology*, Guilan, 90: 141-146, 2003.

SOHNI RY, KAIMAL P, BHATT MR. The antiamoebic effect of a crude drug formulation of herbal extracts against *Entamoeba histolytica* *in vitro* and *in vivo*. *Journal Ethnopharmacology*, Bombay, 45: 43-52, 2003.

STEVENS, K. R.; MYLECRANE, L. Issues in chronic toxicology. In: HAYES, A. W. *Principles and Methods of Toxicology*. 3. ed. New York: Raven Press, 1994. Chapter 18, p. 673-695

SHUBERT S., ANDRESEN B.H., BAHR L., FISCHER L., STAMP R., STRICK G., WITTKE J. W., ULLMANN U. 1996. The immunomodulatory effects of antibiotics: *in vivo* investigations of 21 substances by means of lymphocyte transformation test. *Zentralbl. Bakteriol*, 284: 402-438.

TEO, S., STIRLING, D. THOMAS, S., HOBERMAN, A., KIOPPES, A., KHETANI, V. A 90 day oral gavage toxicity study of D- methylphenidate and D, L-methylphenidate in Sprague Dawley rats. *Toxicology*, Ireland, v. 179, p. 183-196, 2002.

THE WORLD HEALTH REPORT 1996. Fighting Disease, Fostering Development. Geneva: WHO, 1996.

THOMPSON, R.C.A., LYMBERY, A.J., MELONI, B.P. Gentic Variation in giardia. *Kunstler, Protozoological Abstracts*, 14: 1-28, 1990

TOWNSON, S.M., BOREHAM, P.F.,UPCROFT, J.A.. Resistance to the Nitroheterocyclic drugs. *Acta Trop.*, Australia, 56: 173-194, 1990.

UNGAR, B.L.P., YOLKEN, R.H., NASH, T.E., QUINN, T.C. Enyme-linked immunosorbent assay for the detection of *Giardia lamblia* in fecal specimens. *Journal of Infectious Diseases*, 149: 90-97, 1984.

UNITED NATIONS. *The Globally Harmonized System of Classification and Labelling of Chemicals*: New York and Genebra, 2005

VAN DEN HEUVEL, M.J.; CLARCK, D.G.; FIELDER, R.J.; KOUNDAKJIAN, P.P.; OLIVER, G.J.A.; PELLING, D.; TOMLINSON, N.J.; WALKER, A.P. The international validation of fixed dose procedure as an alterantive to the classical LD₅₀ test. *Food and Chemical Toxicology*, England, v. 28, n. 7,p. 469-482, 1990.

YAM, J.; REER, P. J.; BRUCE, R. D. Comparison of the up-and-down method and the fixed-dose procedure for acute oral toxicity testing. *Food and Chemical Toxicology*, England, v. 29, n. 4, p. 259-263, 1991.

YUAN G., GONG Z., LIJ., LI X.. Ginkgo Biloba Extract Protects Against alcohol- induced liver Injury in Rats.*Phytotherapy Research, in press*, 2006

ZIMMERMANN M. Ethical guidelines for investigations of Experimental pain in conscious animals. *Acta Physiologica Scandinavica Supplement*, England, **554**: 221–233, 1986.