Category Justification – The University of Texas at San Antonio

Guinea pigs challenged with *Chlamydia* spp. (*C. caviae*, *C. trachomatis*) develop an infection localized to the genital epithelium and do not experience pain or distress (as evidenced by lack of overt signs of illness) during resolution of the infection (approximately 36 days from time of challenge). This is similar to the situation in human females where approximately 80% are asymptomatic. Additionally, in the available literature, there have been no reports of Guinea Pigs displaying morbidity or mortality after genital chlamydial infection. However, the genital chlamydial infection may result in upper genital tract sequelae, such as infertility.

There is a lack of evidence in the literature that animals infected with this organism experience pain and distress. Additionally, the use of pharmacological agents such as analgesics will alter the immune response and there is significant evidence in the scientific literature showing that the administration of pain medication interferes with immune functions (for example with antigen processing and presentation; the inflammatory mechanisms involved in the initiation; and cytokine differentiation and recruitment of T cells to the joints or other sites of inflammation; as has been shown for NSAID's (Refs: 1-6), morphines (opioids) (Refs: 7-8) or steroids (Refs: 9-12). Moreover, it is not predictable how specific drugs for pain control may affect the expected outcome of our studies. It is likely that pain control will alter both T cell and APC function and will jeopardize the outcome of our study.

- 1. Crotty, B., P. Hoang, H. R. Dalton, and D. P. Jewell. 1992. Salicylates used in inflammatory bowel disease and colchicine impair interferon-gamma induced HLA-DR expression. Gut 33:59.
- 2. Schleimer, R. P., and E. Benjamini. 1981. Effects of prostaglandin synthesis inhibition on the immune response. Immunopharmacology 3:205.
- 3. Neal, T. M., M. C. Vissers, and C. C. Winterbourn. 1987. Inhibition by nonsteroidal anti-inflammatory drugs of superoxide production and granule enzyme release by polymorphonuclear leukocytes stimulated with immune complexes or formyl-methionyl-leucyl-phenylalanine. Biochem. Pharmacol. 36:2511.
- 4. Panush, R. S., and C. R. Anthony. 1976. Effects of acetylsalicylic acid on normal human peripheral blood lymphocytes. Inhibition of mitogen- and antigen-stimulated incorporation of tritiated thymidine. Clin. Exp. Immunol. 23:114.
- 5. Kang, B. S., E. Y. Chung, Y. P. Yun, M. K. Lee, Y. R. Lee, K. S. Lee, K. R. Min, and Y. Kim. 2001. Inhibitory effects of anti-inflammatory drugs on interleukin-6 bioactivity. Biol. Pharm. Bull. 24:701.
- 6. Barasoain, I., J. M. Rojo, and A. Portoles. 1980. "In vivo" effects of acetylsalicylic acid and two ether derived compounds on primary immune response and lymphoblastic transformation. Immunopharmacology 2:293.
- 7. Ni, X., K. R. Gritman, T. K. Eisenstein, M. W. Adler, K. E. Arfors, and R. F. Tuma. 2000. Morphine attenuates leukocyte/endothelial interactions. Microvasc. Res. 60:121.
- 8. Piersma, F. E., M. A. Daemen, A. E. Bogaard, and W. A. Buurman. 1999. Interference of pain control employing opioids in in vivo immunological experiments. Lab Anim 33:328.
- 9. Angeli, A., R. G. Masera, M. L. Sartori, N. Fortunati, S. Racca, A. Dovio, A. Staurenghi, and R. Frairia. 1999. Modulation by cytokines of glucocorticoid action. Ann. N. Y. Acad. Sci. 876:210.
- 10. Ashwell, J. D., F. W. Lu, and M. S. Vacchio. 2000. Glucocorticoids in T cell development and function*. Annu. Rev. Immunol. 18:309.
- 11. Cidlowski, J. A., K. L. King, R. B. Evans-Storms, J. W. Montague, C. D. Bortner, and F. M. Hughes, Jr. 1996. The biochemistry and molecular biology of glucocorticoid-induced apoptosis in the immune system. Recent Prog. Horm. Res. 51:457.
- 12. Nieto, M. A., and A. Lopez-Rivas. 1992. Glucocorticoids activate a suicide program in mature T lymphocytes: protective action of interleukin-2. Ann. N. Y. Acad. Sci. 650:115.