



Harderian Gland – Inflammation



Figure Legend: Figure 1 Harderian gland - Inflammation, Chronic active in a female Sprague- Dawley rat from a chronic study. Numerous inflammatory cells are present in the interstitium and alveoli with destruction of the acinar tissue. Figure 2 Harderian gland - Inflammation, Chronic active in a female Sprague- Dawley rat from a chronic study (higher magnification of Figure 1). The inflammatory cells, largely neutrophils and macrophages expand the interstitium and infiltrate the alveoli, many of which are necrotic. Figure 3 Harderian gland - Inflammation, Suppurative in a male B6C3F1 mouse from a chronic study. There are massive accumulations of neutrophils (arrow) with necrotic debris (asterisk) in the alveolar lumens and interstitium with destruction of the normal architecture. Figure 4 Harderian gland - Inflammation, Chronic study. There are



Harderian Gland – Inflammation



macrophages and lymphocytes (predominantly). interstitial fibrosis, and foreign bodies (hair shaft fragments) (arrow) in the Harderian gland.

Comment: Harderian gland inflammation can be secondary to trauma from retrobulbar bleeding procedures and can be result from other causes, such as infectious agents (e.g., sialodacryoadenitis virus), foreign bodies, excessive exposure to light, and certain nutritional deficiencies.

In NTP studies, there are five standard categories of inflammation: acute, suppurative, chronic, chronicactive, and granulomatous. In acute inflammation, the predominant infiltrating cell is the neutrophil, though fewer macrophages and lymphocytes may also be present. There may also be evidence of edema or hyperemia. The neutrophil is also the predominant infiltrating cell type in suppurative inflammation, however, in suppurative inflammation, the neutrophils are aggregated and many of them are degenerate (suppurative exudate). Cell debris, both from the resident cell populations and infiltrating leukocytes, proteinaceous fluid containing fibrin, fewer macrophages, occasional lymphocytes or plasma cells, and, possibly, an infectious agent may also be present in within the exudate. Grossly, these lesions would be characterized by the presence of pus. In the tissue surrounding the exudate, there may be fibroblasts, fibrous connective tissue, and mixed inflammatory cells, depending on the chronicity of the lesion. Lymphocytes predominate in chronic inflammation. Lymphocytes also predominate in chronic-active inflammation, but in chronic-active inflammation, there are also a significant number of neutrophils. Both lesions may contain macrophages. Granulomatous inflammation is another form of chronic inflammation, but this diagnosis requires the presence of a significant number of aggregated, large, activated macrophages, epithelioid macrophages, or multinucleated giant cells. Inflammation differs from cellular infiltration in that inflammatory lesions have evidence of tissue damage, such as edema, hemorrhage, degeneration, necrosis, regeneration, etc.

Recommendation: Harderian gland inflammation should be diagnosed and assigned a severity grade. An appropriate type modifiers (acute, chronic, etc.) should be included in the diangosis. Foreign bodies in the Harderian gland can be diagnosed separately as present (without assignment of a severity grade). Associated lesions (e.g., necrosis) should not be diagnosed separately (unless warranted by severity), though they can be described in the narrative.





Harderian Gland – Inflammation

References:

Beaumont SD. 2002. Ocular disorders of pet mice and rats. Vet Clin Exot Anim 5:311-324. Abstract: <u>http://www.ncbi.nlm.nih.gov/pubmed/12170635</u>

Botts S, Jokinen M, Gaillard ET, Elwell MR, Mann PC. 1999. Salivary, Harderian, and lacrimal glands. In: Pathology of the Mouse: Reference and Atlas (Maronpot RR, Boorman GA, Gaul BW, eds). Cache River Press, Vienna, IL, 49-79.

Abstract: http://www.cacheriverpress.com/books/pathmouse.htm

Greaves P. 2007. Nervous system and special sense Organs. In: Histopathology of Preclinical Toxicity Studies: Interpretation and Relevance in Drug Safety Evaluation, 3rd ed. Academic Press, San Diego, CA, 861-933.

Abstract: http://www.sciencedirect.com/science/book/9780444527714

Krinke AL, Schaetti PR, Krinke GJ. 1994. Changes in the major ocular glands. In: Pathobiology of the Aging Rat, Vol 1 (Mohr U, Dungworth DL, Capen CC, Carlton WW, Sundberg JP, Ward JM, eds). International Life Sciences Institute Press, Washington, DC, 109-119.

Krinke GJ, Schaetti PR, Krinke A. 1996. Nonneoplastic and neoplastic changes in the Harderian and lacrimal glands. In: Pathobiology of the Aging Mouse, Vol 2 (Mohr U, Dungworth DL, Capen CC, Carlton WW, Sundberg JP, Ward JM, eds). International Life Sciences Institute Press, Washington, DC, 139-152.

Lambert RA, Yudkin AM. 1923. Changes in the paraocular glands accompanying the ocular lesions which result from a deficiency of vitamine A. J Exp Med 38:25-32. Abstract: http://www.ncbi.nlm.nih.gov/pubmed/19868768

National Toxicology Program. 1989. NTP TR-345. Toxicology and Carcinogenesis Studies of Roxarsone (CAS No. 121-19-7) in F344/N Rats and B6C3F₁ Mice (Feed Studies). NTP, Research Triangle Park, NC.

Abstract: http://ntp.niehs.nih.gov/go/8644

National Toxicology Program. 1989. NTP TR-350. Toxicology and Carcinogenesis Studies of Tribromomethane (Bromoform) (CAS No. 75-25-2) in F344/N Rats and B6C3F₁ Mice (Gavage Studies). NTP, Research Triangle Park, NC. Abstract: http://ntp.niehs.nih.gov/go/6961

National Toxicology Program. 2007. NTP TR-526. Toxicology and Carcinogenesis Studies of a Mixture of 2,3,7,8-Tetrachlorodibenzo-p-Dioxin (TCDD) (CAS No. 1746-01-6), 2,3,4,7,8-Pentachlorodibenzofuran (PeCDF) (CAS No. 57117-31-4), and 3,3',4,4',5-Pentachlorobiphenyl (PCB 126) (CAS No. 57465-28-8) in Female Harlan Sprague-Dawley Rats (Gavage Studies). NTP, Research Triangle Park, NC.

Abstract: http://ntp.niehs.nih.gov/go/9307





Harderian Gland – Inflammation

References:

O'Steen WK, Kraeer SL, Shear CR. 1978. Extraocular muscle and Harderian gland degeneration and regeneration after exposure of rats to continuous fluorescent illumination. Invest Ophthalmol Vis Sci 17:847-856.

Abstract: http://www.iovs.org/content/17/9/847.short

Payne AP. 1994. The Harderian gland: A tercentennial review. J Anat 185:1-49. Abstract: <u>http://www.ncbi.nlm.nih.gov/pmc/articles/PMC1166813/</u>

Percy DH, Wojcinski ZW, Schunk MK. 1989. Sequential changes in the Harderian and exorbital lacrimal glands in Wistar rats infected with sialodacryoadenitis virus. Vet Pathol 26:238-245. Full-text: <u>http://vet.sagepub.com/content/26/3/238.full.pdf</u>

Rothwell TLW, Everitt AV. 1986. Exophthalmos in ageing rats with Harderian gland disease. Lab Anim 20:97-100. Abstract: http://lan.sagepub.com/content/20/2/97.short

Author:

Margarita M. Gruebbel, DVM, PhD, DACVP Senior Pathologist Experimental Pathology Laboratories, Inc. Research Triangle Park, NC