

NTP Nonneoplastic Lesion Atlas

Testis, Germ cell – Exfoliation

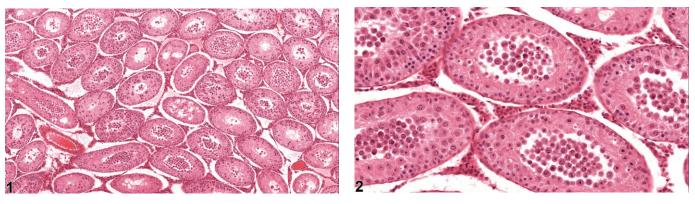


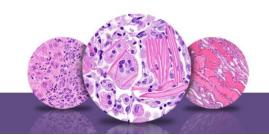
Figure Legend: Figure 1 Testis, Germ cell - Exfoliation in a male Harlan Sprague-Dawley rat. Exfoliated germ cells are present in seminiferous tubule lumens. (Photograph courtesy of Dr. D. Creasy.) **Figure 2** Testis, Germ cell - Exfoliation in a male Harlan Sprague-Dawley rat. Higher magnification of Figure 1 showing germ cells in seminiferous tubule lumens. (Photograph courtesy of Dr. D. Creasy.)

Comment: Germ cells may undergo degeneration and/or apoptosis within the seminiferous epithelium, or they may lose contact with the surrounding Sertoli cell cytoplasmic processes and be shed into the tubular lumen (Figure 1 and Figure 2). Often the exfoliated germ cells retain relatively normal morphology. The cells will rapidly be transported into the epididymis and will be present in the ductular lumens. The finding should not be confused with artifactual sloughing of germ cells caused by squeezing of the testis during necropsy dissection. In the case of sloughing due to necropsy trauma, germ cells will not be seen in the epididymis. Germ cell exfoliation has been described as a predominant finding following administration of phthalate esters and microtubule inhibitors such as colchicine and the fungicide carbendazim.

Recommendation: The term "germ cell exfoliation" should be reserved for those situations where there is extensive shedding of germ cells into the lumen as a predominant finding. In those situations, germ cell exfoliation should be diagnosed and graded and should be discussed in the pathology narrative if the incidence and/or severity appears to be related to chemical administration. Bilateral involvement should be diagnosed when present. If there is significant concurrent germ cell degeneration, that should also be diagnosed and given a severity grade.



NTP Nonneoplastic Lesion Atlas



Testis, Germ cell – Exfoliation

References:

Correa LM, Nakai M, Strandgaard CS, Hess RA, Miller MG. 2002. Microtubules of the mouse testis exhibit differential sensitivity to the microtubule disruptors carbendazim and colchicine. Toxicol Sci 69:175-182.

Abstract: http://www.ncbi.nlm.nih.gov/pubmed/12215672

Creasy DM. 2001. Pathogenesis of male reproductive toxicity. Toxicol Pathol 29:64-76. Full Text: <u>http://tpx.sagepub.com/content/29/1/64.full.pdf</u>

Creasy D, Bube A, de Rijk E, Kandori H, Kuwahara M, Masson R, Nolte T, Reams R, Regan K, Rehm S, Rogerson P, Whitney K. 2012. Proliferative and nonproliferative lesions of the rat and mouse male reproductive system. Toxicol Pathol 40:40S-121S. Abstract: http://www.ncbi.nlm.nih.gov/pubmed/22949412

Nakai M, Hess RA. 1994 Morphological changes in the rat Sertoli cell induced by the microtubule poison carbendazim. Tissue Cell 26:917-127. Abstract: <u>http://www.ncbi.nlm.nih.gov/pubmed/7886678</u>

Russell LD, Malone JP, MacCurdy DS. 1981. Effect of the microtubule disrupting agents, colchicine and vinblastine, on seminiferous tubule structure in the rat. Tissue Cell 13:349-367. Abstract: <u>http://www.ncbi.nlm.nih.gov/pubmed/7314074</u>

Authors:

Dianne M. Creasy, PhD, Dip RCPath, FRCPath Dianne Creasy Consulting LLC Pipersville, PA

Robert R. Maronpot, DVM, MS, MPH, DACVP, DABT, FIATP Senior Pathologist Experimental Pathology Laboratories, Inc. Research Triangle Park, NC

Dipak K. Giri, DVM, PhD, DACVP Toxicologic Pathologist Integrated Laboratory Systems, Inc. Research Triangle Park, NC