# THE HISTOLOGY OF THE URINARY SYSTEM OF THE MINK, MUSTELA VISION

Thesis for the Degree of M. S. MICHIGAN STATE UNIVERSITY Richard Edward Bostrom 1966





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# THE HISTOLOGY OF THE URINARY SYSTEM

#### OF THE MINK, MUSTELA VISION

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Richard Edward Bostrom

#### A THESIS

Submitted to Michigan State University in partial fulfillment of the requirements for the degree of

MASTER OF SCIENCE

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ii

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### TABLE OF CONTENTS

																									Page
ACKNOW	VLED	GME	NI	'S	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	ii
LIST C	OF I	ABL	ES	5	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	vi
LIST C	OF F	FIGU	JRE	S	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	vii
INTROL	DUCI	TON	I	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	1
MATERI	EALS	S AN	ID	ME	TH	101	S	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	2
RESULI	rs A	ND	DI	sc	CUS	SI	101	1	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	6
Kić	lney	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	6
	Cap Uri	sul nif	.e 'er	• •00	IS	Tu	ibi	ile	es	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	6 6
		Nep	hr	or	ı		•	•	•	•	•	•	•	•			•	•	•	•	•	•	•	•	6
			Re Pr Lc Di	ena ox op	il in o c al	Cc al of . C	org L ( He Cor	ous Cor en]	sci nvo Le	le blu	ite ed	ed Tu	Tu Ibu	ibu ile	ile	•	• • •	•	• • •	• • •	• • •	• • •	• • •	• • •	6 7 8 8
		Col	.le	ect	;ir	ng	Τι	ıbı	ile	es	•				•					•				•	9
			Ar Pa	ch ipi	neć	l a .ar	ind 'y	i S Di	Sti ici	rai ts	gł	nt •	Cc	)]] •	.ec	ti.	ng •	; [	uc	ts •	·	•	•	•	9 9
	Mac Jux	ula tag	L D C	)er	isa eru	ı ıla	ır	Aŗ	ppa	ara	itu	IS	•	•	•	•	•	•	•	•	•	•	•	•	10 10
Ure	eter	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	11
	Tur	nica	ιM	luc	eos	sa	•	•	•	•	•	•	•	•			•	•	•	•	•	•	•	•	12
		Epi Lam	.th ir	nel na	iu Pr	im 'or	ori	La	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	12 12
	Tur Tur	nica nica	L M L A	lus Idv	scı ver	ila iti	ari Lti	is ia	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	13 13

•

ï

. . .

# Page

	Uri	nar	уB	ladd	ler	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	13
	,	Tun	ica	. Muc	cosa	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	13
			Epi	.the]	Lium	•	•	•	•	•	•	•	•	•	•	•	•	•	•		•	•	13
			Lam	ina	Pro	pr	ia	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	15
			Mus	cula	aris	Mı	uco	osa	аe	•	•	•	•	•	•	•	•	•	•	•	•	•	15
	ı	Tun	ica	Sut	omuc	os	a	•			•	•	•	•	•	•		•	•		•	•	15
	'	Tun	ica	Mus	scul	ar	is	•	•	•		•	•	•	•		•	•	•	•	•	•	15
	1	Tun	ica	. Adv	vent	it	ia	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	16
	Fem	ale	Ur	ethr	ra .	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	16
	ı	Tun	ica	Muc	cosa	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	16
			Epi	the]	<b>1 1</b> ນຫ													-					16
			Lam	ina	Pro	pr	ia															•	17
						<b>F</b> = 1							-		-	-	-	-	-	-	-	-	- •
	Mal	e U	ret	hra	• •	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	17
	ı	Tun	ica	Muc	cosa	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	17
			Epi	the]	Lium																		17
			Lam	ina	Pro	pr	ia	•	•		•	•	•	•	•	•	•		•	•	•		18
						•																	
SUM	MAR	Υ.	•	•••	• •	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	19
LIT	ERA	TUR	ЕC	ITEI	ο.	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	22

## LIST OF TABLES AND ILLUSTRATIONS

Table		Page
1.	Specimens used in the study	3
2.	Analysis of the normal ranch diet fed the animals used in this study	3
3.	Summary of kidney measurements	11
Illustrat	ions	

1.	Areas of	the ur	inary s	ystem	from whi	lch				
	tissues	were	selected	d for	study	•	•	•	•	5

### LIST OF FIGURES

Figure		Page
1.	Kidney capsule	25
2.	Reticular network of the kidney and its capsule	27
3.	Kidney cortex	29
4.	Kidney cortex	29
5.	Branching arcuate artery near the cortico- medullary junction	31
6.	Concentrations of lipid material in the proximal convoluted tubules and the thick descending limbs of Henle	33
7.	Lipid droplets concentrated in the proxi- mal convoluted tubules and the thick descending limbs of Henle	35
8.	P.A.S. positive material in the epithelium covering the renal crest	35
9.	P.A.S. positive material in the epithelium lining the papillary ducts	37
10.	P.A.S. positive material in the epithelium lining the papillary ducts	37
11.	Proximal ureter with surrounding fat tunic	39
12.	Dense reticular network in the lamina pro- pria and tunica submucosa of the ureter	41
13.	Arrangement of the muscular coats in the tunica muscularis of the proximal ureter	43
14.	Cross section of the middle of the ureter	45
15.	Cross section of the distal ureter	47
16.	Transitional epithelium of the ureter	49

vii

# Figure

17.	Transitional epithelium of the ureter	51
18.	Reticular network of the bladder	53
19.	Arrangement of the tunica muscularis of the bladder	53
20.	Urinary bladder mucosa	55
21.	Transitional epithelium of the bladder	55
22.	Bladder epithelium with surface cells about to be sloughed	57
23.	Cross section of the outer bladder wall .	59
24.	Cross section of the prostatic urethra	59
25.	Prostatic duct opening into the urethral lumen	61
26.	Cavernous spaces in the prostatic urethra	63
27.	Elastic fibers in the lamina propria of the prostatic urethra	63
28.	Prostatic utriculus	65
29.	Cross section of the membranous urethra .	67
30.	Mucosa of the membranous urethra	67
31.	Epithelial lining of the membranous urethra	69
32.	Cross section of the membranous urethra showing the extensive network of elastic fibers	71
33.	Cross section of the membranous urethra showing the extensive network of elastic fibers	71
34.	Cross section of the proximal penile urethral mucosa	73
35.	Epithelial lining of the proximal penile urethra	73
36.	Cross section of the penile urethra near the external urethral orifice	75

# Figure

37.	Epithelial lining of the penile urethra near the extrenal urethral orifice	75
38.	Reticular network of the female urethra .	77
39.	Cross section of the proximal female urethra	77
40.	Female urethral epithelium	79
41.	Nests of serous glands in the submucosa and between the bands of the skeletal urethral muscle in the female urethra .	79
42.	Intraepithelial cysts in the female urethral epithelium	81
43.	Epithelial lining of the distal female urethra	81

Page

#### INTRODUCTION

From an economic standpoint, losses suffered due to urinary tract pathology such as urolithiasis (Nielson, 1956), urinary incontinence (Schaible <u>et al</u>., 1962; Aulerich <u>et al</u>., 1963), "Wet Belly" (Leoschke, 1962; Aulerich <u>et al</u>., 1963), and Aleutian disease (Karstad and Pridham, 1962; Leader <u>et al</u>., 1963; Gershbein and Spencer, 1964; Thompson and Aliferis, 1964) are of extreme importance to the mink rancher. A thorough review of the literature indicated that there were no reports available on the normal histology of the urinary system of the mink. Because a previous knowledge of the normal is essential in recognizing the abnormal, a histological study of the urinary tract of the normal mink (Mustela vision) was undertaken.

#### MATERIALS AND METHODS

The entire urinary systems of eight male and eight female clinically normal mink were supplied by the Michigan State University mink ranch (Table 1). The animals, ranging in age from one month to two years, were fed a normal ranch diet (Table 2).

Twelve of the animals were killed by cervical dislocation. Their urinary tracts were dissected out and immediately placed in a 10% formalin solution buffered with sodium acetate or in Carnoy's fixative (Gridley, 1957). The tissues were kept in the fixative for 48 hours prior to embedding.

Because of the extreme difficulty in preserving the cytological detail of the kidney, the urinary systems of the remaining four animals were fixed with a perfusion of buffered formalin. This was accomplished by first anesthetizing the animals with an intraperitoneal injection of 3% sodium pentabarbital (1 ml./kg. of body weight). After the animals were anesthetized, the carotid artery and the external jugular vein were cannulated and the vascular system flushed via the carotid artery with mammalian Ringer's solution. This was followed immediately with an infusion of buffered formalin. After infusion, the urinary tract was dissected out and placed in a solution of buffered formalin for 48 hours.

Specimen	Mink	Age in	Weight	Sex	Color		
Number	Number	Months	in Grams		Pattern		
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16	K 381 KP 181 K 371 KP 2 KP 52 K 264 K 470 L 70 KP 183 JS 322 K 310 KP 707 LP 272 L 167 K 315 L 631	10 12 14 2 14 1 2.5 14 24 13 12 7 7 12 6	1,550 1,460 1,850 950 480 815 155 705 1,600 770 965 1,800 785 1,300 1,635 965	Male Male Female Female Female Female Female Female Female Male Female Male Male Male Male	Dark Pastel Dark Pastel Pastel Dark Dark Pastel Sapphir Dark Pastel Pastel Dark Dark Dark Dark		

Table 1.--Specimens used in the study.

Table 2.--Analysis of the ration fed to the animals used in this study.

Moisture	•	•	•	•	•	•	•	•	•	•	67.99%
Protein	•	•	•	•	•	•	•	•	•	•	10.81%
Ether extract	•	•	•	•	•	•	•	•	•	•	13.35%
Crude fiber	•	•	•	•	•	•	•	•	•	•	5.69%
Ca	•	•	•	•	•	•	•	•	•	•	.22%
Ph	•	•	•	•	•	•	•	•	•	•	.23%
Na	•	•	•	•	•	•	•	•	•	•	.17%
к	•	•	•	•	•	•	•	•	•	•	.14%
Trace minerals	•	•	•	•	•	•	•	•	•	•	1.40%

Just prior to embedding, the tissues were removed from the fixatives and trimmed to block size. Areas from which tissues were selected for study are shown in Illustration 1. The tissues were dehydrated and cleared in four changes of dioxane (Bucher and Blakely, 1936) and infiltrated with Paraplast under a vacuum of 25 mm./Hg. at 58-60°C. for a period of one to two hours.

Unless otherwise indicated, sections were cut 6-8 microns. Hematoxylin and eosin (Malewitz and Smith modification, 1955), Weigert and Van Gieson's connective tissue stain (Mallory, 1944) and Periodic Acid-Schiff (Gridley, 1957) were used routinely. The following special techniques were also used: (1) A modified reticular tissue stain (Gridley, 1957; Lillie, 1954); (2) Alcian blue and nuclear fast red (Gridley, 1957) for acid mucopolysaccharides; (3) Crossmon's (1937) modification of Mallory's triple stain for connective tissue; and (4) Oil Red O (Mallory, 1944) and Sudan Black B (Chieffele and Putt, 1951) for lipid material. The last two stains were used on frozen sections as most lipids are dissolved by the paraffin embedding technique.





#### RESULTS AND DISCUSSION

#### KIDNEY

The mink kidney is bean-shaped and unipyramidal with the papillary ducts opening on a renal crest rather than a renal papilla. This is similar to the cat and dog kidney (Trautmann and Fiebiger, 1957).

#### CAPSULE

The capsule of the mink kidney is composed of an inner and an outer coat. The outer coat is much thicker and consists almost entirely of adipose tissue with a few blood vessels and nerves coursing through it (Figure 1). The inner coat is primarily fibrous connective tissue interlaced with reticular fibers, some of which penetrate the cortex of the kidney (Figure 1). No smooth muscle or elastic connective tissue is noted. This is similar to the inner one-layered tunica fibrosa of the cat kidney (Yadava and Calhoun, 1958).

#### URINIFEROUS TUBLES

#### Nephron

Renal Corpuscie.--The renal corpuscies as well as the glomeruli are spherical and have an average transverse diameter of 115 and 95.5 microns, respectively. The parietal

and visceral layers of Bowman's capsule, as in other animals, are lined with a layer of simple squamous epithelium. The intercapillary space contains a few reticular fibers and an occasional fibroblast.

Proximal Convoluted Tubule .-- The average transverse diameter and epithelial height of the proximal convoluted tubules are 39.9 and 11.3 microns, respectively. They are lined by a single layer of pyramidal and/or low columnar shaped cells with granular acidophilic cytoplasm (Figure 3). Numerous intracytoplasmic lipid droplets, which are Sudan Black B and Oil Red O positive, are present in the epithelium lining the proximal tubules and those of the thick descending limb of Henle as well (Figures 6 and 7). Similar lipid material has been reported in the cat kidney (Modell, 1933; Lobban, 1955; Stranack, 1962), in the dog kidney (Mac Nider, 1945) and in the kidney of the lion, tiger, cheetah and ocelot (Hewer et al., 1948). In most other animals and man this is considered pathologic. The basal portions of the cells lining the proximal convoluted tubules have a striated appearance due to the presence of numerous mitochondria in the form of rods and filaments which line up perpendicular to the basement membrane. The apical surface of the epithelium has a prominent brush border which increases the surface area of the cells. Its mucopolysaccharide makeup is evidenced by its P.A.S. positive reaction (Figures 4 and 5). The nuclei are large and usually spherical, contain a prominent nucleolus and have a coarse chromatic network. Cell boundaries are indistinct.

Loop of Henle.--The loop of Henle consists of a descending and an ascending portion connected by a sharp bend. It is divided into a thick and a thin segment. The epithelium lining the thick descending portion resembles that of the proximal convoluted tubule, and the epithelium of the thick ascending portion resembles that of the distal convoluted tubule. The thin segment has a diameter of 17.4 microns which is smaller than either the proximal or the distal tubules. It is lined by simple squamous or low cuboidal epithelium. Spherical bulging nuclei give an irregular contour to the lumen ot these tubules (Figure 7). The nuclei have a distinct nuclear membrane, a nucleolus and a dense granular chromatin network. The cytoplasm is slightly eosinophilic. A brush border is not visible with the light microscope.

Distal Convoluted Tubule.--The distal convoluted tubules have an average transverse diameter and epithelial height of 35.6 and 8.0 microns, respectively. They are lined by simple cuboidal epithelium. The cells have fine basal striations with faint borders. They are lower and smaller than those in the proximal tubules, so in a typical cross section more nuclei are seen. Their agranular cytoplasm is slightly more eosinophilic than that of the thin loop of Henle. The nuclei are characterized by a prominent nucleolus, a dense chromatin network, and a distinct nuclear membrane. Lipid material is not evident in the distal convoluted tubules (Figure 7).

Collecting Tubules

Arched and Straight Collecting Ducts.--The arched and straight collecting ducts are difficult to separate histologically and will be discussed together. They have an average transverse diameter and epithelial height of 33.1 and 7.0 microns, respectively. Light, eosinophilic staining, low cuboidal cells with distinct cell boundaries characterize the epithelium lining these ducts. They contain spherical, basally placed nuclei with an indistinct nucleolus and dense chromatin networks that stain darkly with hematoxylin. Basal striations are not apparent, and a brush border is lacking. The cells of the straight collecting tubules contain a few P.A.S. positive, Alcian blue negative, intracytoplasmic granules.

Papillary Ducts.--The papillary ducts have an average transverse diameter and epithelial height of 53.8 and 9.9 microns, respectively. The epithelial lining changes from cuboidal or low columnar at the origin of the ducts to high columnar and sometimes transitional as the ducts open into the renal pelvis. The cells have distinct boundaries but lack basal striations and brush borders. The cytoplasm is colorless or slightly eosinophilic. P.A.S. positive, Alcian blue negative intracytoplasmic granules are present in many of the cells lining these ducts (Figures 9 and 10). They are often most prominent in the apical portion of the cells and may occasionally be seen in the lumen of the tubules. In some cases these granules are found in the cells

of the epithelium covering the renal crest (Figure 8). These granules are similar to those mentioned by Longley <u>et al.</u>, (1963) in the collecting tubules of normal guinea pigs and those in the pelvic epithelium covering the pyramids in the normal human kidney by Tucker <u>et al.</u>, (1959). Karstad (1964) reported similar granules in 23 out of 40 mink showing lesions pathognomonic for Aleutian disease. These granules were also found in 8 out of 14 normal mink from the same study.

#### MACULA DENSA

The macula densa is a modification of that portion of the distal convoluted tubule that comes in contact with the juxtaglomerular apparatus of the afferent arteriole (Figures 3 and 4). This portion is lined by closely packed columnar shaped cells that lack distinct cell boundaries and basal striations. Visible granulation is lacking in the faintly acidophilic cytoplasm. The closely packed spherical or oval nuclei are larger in this area and seem to have a greater affinity for hematoxylin.

#### JUXTAGLOMERULAR APPARATUS

The juxtaglomerular apparatus is located near the vascular pole of the glomerulus where the afferent arteriole comes in contact with the macula densa of the distal convoluted tubule. It consists of enlarged, slightly basophilic, myoepithelioid cells in the tunica media of the afferent

arteriole (Figure 3). A summary of the kidney measurements is shown in Table 3.

Table 3.--Summary of kidney measurements.

1.	Renal corpuscle	115.0
2.	Glomerulus	95.5
3.	Proximal convoluted tubule	
	a. Cross sectional diameter b. Epithelial height	39.9 11.3
4.	Thin loop of Henle	
	<ul><li>a. Cross sectional diameter</li><li>b. Epithelial height</li></ul>	17.4 4.6
5.	Distal convoluted tubule	
	a. Cross sectional diameter b. Epithelial height	35.6 8.0
6.	Arched and straight collecting ducts	
	a. Cross sectional diameter b. Epithelial height	33.1 7.0
7.	Papillary duct	
	a. Cross sectional diameter b. Epithelial height	53.8 9.9

#### URETER

The wall of the ureter is generally divided into tunica mucosa, tunica submucosa, tunica muscularis, and tunica adventitia. In the following discussion, the mucosa and submucosa will be described together as there is no sharp demarcation between the two.

Microns

#### TUNICA MUCOSA

#### Epithelium

The epithelium is transitional and varies from 1 to 8 cell layers in thickness (Figures 16 and 17). It has a cross sectional diameter varying from 9 to 98 microns. This wide variation in cell layers and cross sectional diameters is primarily due to the different degrees of lumenal distention at the time of fixation. Elongated spindle-shaped cells with hyperchromatic nuclei and slightly more eosinophilic cytoplasm than the remainder of the epithelium could occasionally be seen on the luminal border of the epithelium (Figures 15 and 16). P.A.S. positive material similar to that reported by Liu (1962) is present in the epithelial cytoplasm.

#### Lamina Propria

The lamina propria consists of loosely arranged collagenous connective tissue, blood vessels, lymphatics, nerves, diffuse lymphatic tissue, a few elastic fibers, and a supporting network of reticular fibers (Figure 12). The loose arrangement allows the epithelium to be thrown into longitudinal folds, and this folding is responsible for the characteristic stellate appearance of the lumen (Figure 15). There is an average of 6 major and minor folds in the proximal segment of the ureter, 6 in the middle segment and 5 in the distal segment of the ureter. A capillary plexus is present just beneath the epithelium (Figure 16). This is

similar to the capillary plexus present in the ureters of ruminants (Calhoun, 1959).

#### TUNICA MUSCULARIS

The tunica muscularis of the ureter consists of an inner longitudinal, middle circular, and an outer longitudinal layer of smooth muscle. The inner longitudinal layer thins out and the outer longitudinal layer thickens near the bladder (Figures 13, 14, and 15). The middle circular layer maintains a relatively constant thickness the entire length of the ureter (Figures, 13, 14, and 15). The dog, cat, pig, cow, sheep, goat and horse have a similar muscular arrangement (Calhoun, 1959).

#### TUNICA ADVENTITIA

A loosely arranged collagenous connective tissue rich in blood vessels, lymphatics, nerves and containing a few autonomic ganglia is characteristic of the adventitia throughout the length of the ureter (Figure 14). This is surrounded by a thick tunic of fat which is supplied by a sparse vascular and nerve supply (Figure 11).

#### URINARY BLADDER

TUNICA MUCOSA

#### Epithelium

The transitional epithelium lining the urinary bladder varies from three to four cell layers in thickness and from 29 to 46 microns in cross sectional diameter. As in the ureter, this wide variation is primarily due to the different degrees of luminal distention at the time of fixation. The surface epithelial cells often have a slightly, greater affinity for eosin than the rest of the epithelium, and this tends to give a cuticle-like appearance to the luminal border. The large, umbrella, cuboidal and pearshaped cells have oval or spherical vesicular nuclei. They contain one or more nucleoli and have little affinity for hematoxylin (Figures 21 and 22). Occasional elongated spindle-shaped cells with hyperchromatic nuclei are also The basal and intermediary layers consist of cuboidal seen. and columnal cells with spherical or oval nuclei that have a thin nuclear membrane and a faint diffuse chromatin network. The cell boundaries are indistinct and the cytoplasm is slightly eosinophilic. In a number of the areas, the surface cells are elongated and resemble the cornified cells in the stratum corneum of stratified squamous epithelium. In these areas and others, there seem to be a great many cells undergoing degenerative changes. Many of these cells have sloughed into the bladder lumen (Figure 22). This might possibly be a factor in the high incidence of urinary calculi in mink as these sloughed cells could easily act as foci for the deposition of the various mineral salts making up the calculi.

Lamina Propria

The lamina propria consists of a framework of reticular (Figure 18) and dense fibroelastic connective tissue interspersed with scattered lymphocytes, occasional solitary lymphoid nodules, numerous lymphatic and blood vessels and a few nerves. The blood vessels form a rich vascular bed just under the epithelium and occasionally appear to penetrate its basal layer (Figures 20 and 21).

Muscularis Mucosae

Small scattered bundles of smooth muscle arranged in circular, longitudinal and oblique planes comprise the muscularis mucosae. It serves to separate the mucosa from the submucosa and probably helps account for the marked folding of the mucosa in the contracted bladder (Figure 19).

TUNICA SUBMUCOSA

Except for being somewhat thinner, loosely arranged and slightly less vascular, the submucosa is similar to the lamina propria.

#### TUNICA MUSCULARIS

The tunica muscularis consists of two thick layers of smooth muscle usually arranged in an inner circular and outer longitudinal manner (Figure 19). On occasion these layers seem to either spiral or cross and give the appearance of an inner circular, outer longitudinal layer in one place and an inner longitudinal, outer circular layer in

another area on the same section. The muscle layers are supplied by numerous blood vessels, lymphatics and nerves. A few intramural autonomic ganglia are present between the various muscle bundles (Figure 23).

#### TUNICA ADVENTITIA

Loosely and densely arranged collagenous connective tissue, a few elastic fibers, numerous blood vessels, lymphatics, nerves and a few autonomic ganglia constitute the tunica adventitia. In some areas it is covered by a reflection of the peritoneum and, therefore, could be called a serosa (Figure 23).

#### FEMALE URETHRA

TUNICA MUCOSA

#### Epithelium

The lining of the female urethra varies from predominantly transitional epithelium at its origin, mixtures of transitional, stratified and pseudostratified columnar, and stratified squamous in the mid-portions (Figure 40) to primarily stratified squamous at the external urethral orifice (Figure 43). A few intraepithelial nests of mucous cells are present in all segments and are especially numerous in the proximal and middle portions (Figure 40). Invaginations of the epithelium (lacunae of Morgagni) containing saccular outpocketings of mucous secreting cells penetrate the lamina propria and occasionally extend deeply into the
submucosa (Figure 42). The mucous cells in both the intraepithelial nests and lacunae of Morgagni contain P.A.S. positive, Alcian blue negative, secretion granules located primarily in the apical portion of the cytoplasm. In some cases P.A.S. positive eosinophilic concretions resembling intraepithelial cysts could be seen distending the lumens of the tubules and glandular nests (Figure 42).

## Lamina Propria

Fibroelastic and reticular connective tissue penetrated by invaginations of the epithelium from the luminal surface and ducts from the serous intramural and extramural glands in the tunica muscularis surround the numerous venous sinuses of the lamina propria-submucosa (Figures 38, 39, and 41). The venous sinuses are most prominent in the proximal and midportions and diminish somewhat near the urethral orifice.

## MALE URETHRA

TUNICA MUCOSA

#### Epithelium

The epithelium lining the origin of the prostatic urethra is two to four cell layers in thickness, primarily transitional in nature with some areas resembling pseudostratified columnar and stratified cuboidal epithelium. The large spherical to oval nuclei have a medium to heavy chromatin network, a prominent nucleolus, and a heavy

nuclear membrane. Cell boundaries are faint or indistinct. In most cases the basal cells are similar to those on the apical surface. However, a few of the surface cells are noted undergoing degenerative changes, suggesting that they are about to be sloughed.

The prostatic portion of the proximal urethra is surrounded by the prostate gland which consists of multilobular mucous acini which empty via their ducts into the urethral lumen (Figures 24 and 25). The prostatic utriculus and the colliculus seminalis are evident in the middle of this area (Figure 28).

The epithelium lining the membranous and penile urethra is similar to that in the proximal urethra except there is a greater proportion of pseudostratified columnar and stratified columnar epithelium and a decrease in the amount of typical transitional epithelium (Figures 31 and 35). Near the urethral orifice the epithelium is lower and resembles stratified cuboidal epithelium (Figure 37).

Lamina Propria

The lamina propria of the male urethra consists of numerous venous sinuses surrounded by collagenous and reticular connective tissue. The venous sinuses of the prostatic and membranous urethra are less extensive than those in the penile urethra (Figures 26, 29, 34 and 36). In addition, the lamina propria of the male urethra contains numerous elastic fibers throughout its length (Figures 27, 32 and 33).

# SUMMARY

Histologic studies were made on the entire urinary systems of eight male and eight female mink ranging in age from one month to two years.

The mink kidney is unipyramidal. The capsule has a single layered tunica fibrosa. The renal corpuscles and glomeruli are spherical and have an average transverse diameter of 115 and 95.5 microns, respectively. The epithelium of the proximal convoluted tubules and thick descending limbs of Henle have an average transverse diameter and epithelial height of 39.9 and 11.3 microns respectively and contain numerous intracytoplasmic lipid droplets. A brush border is uniformly arranged along the luminal border of these cells. The macula densa and juxtaglomerular apparatus are similar to those of other species.

The epithelium of the straight collecting ducts and papillary ducts contain varying numbers of P.A.S. positive, Alcian blue negative, intracytoplasmic granules which are often located in the apical portion of the cells.

The ureter is lined by transitional epithelium. A capillary plexus is located in the lamina propria adjacent to the base of the epithelium. The tunica muscularis consists of an inner logitudinal, middle circular and outer

longitudinal layer of smooth muscle. The inner layer thins out and the outer layer thickens near the bladder. The tunica adventitia is surrounded by a thick tunic of fat.

The urinary bladder is lined by transitional epithelium of varying thickness. A number of the surface cells have undergone degenerative changes and have sloughed into the bladder lumen. Numerous blood vessels form a rich vascular plexus just below the epithelium. Ossasionally some of these vessels could be seen penetrating the basal layers of the epithelium. Small, scattered, variously arranged bundles of smooth muscle comprise the muscularis mucosae. The tunica muscularis consists of two thick spiraling layers of smooth muscle.

The epithelium lining the female urethra varies from transitional to stratified squamous. Intraepithelial nests of mucous cells, lacunae of Morgagni and intraepithelial cysts are present in variable numbers. The lamina propria-submucosa consists of numerous venous sinuses surrounded by fibroelastic and reticular connective tissue. The venous sinuses are most prominent in the proximal and midportions of the urethra.

The epithelial lining of the male urethra is variable throughout its length. It is predominantly transitional at its origin, variably transitional, stratified and pseudostratified columnar throughout the remainder, except near the urethral orifice where it resembles stratified cuboidal.

Other than the size differences between the very young and the adult animals and the differences between the male and female urethra, no obvious sex or age differences are noted.

## LITERATURE CITED

- Aulerich, R. J., Ringer, R. K., and Schaible, P. J.: Mink pelts affected with "Wet Belly." Quar. Bull. of Mich. Ag. Exp. Sta., 44, (1962):384-391.
- Aulerich, R. J., Schaible, P. J., and Ringer, R. K.: Effect of time of pelting on the incidence of "Wet Belly." Quar. Bull. of Mich. Ag. Exp. Sta., 45, (1963): 439-443.
- Aulerich, R. J., Shelts, S., and Schaible, P. J.: Influence of dietary calcium levels on the incidence of urinary incontinence and "Wet Belly" in mink. Quar. Bull. of Mich. Ag. Exp. Sta., 45, (1963): 444-449.
- Bucher, C. J., and Blakely, K. D.: The use of dioxane in the preparation of histologic sections by the paraffin method. Anat. Rec., 132, (1936):569-583.
- Calhoun, M. L: Comparative histology of the ureters of domestic animals. Anat. Rec., 133, (1959):365-366 (abstract).
- Chiffelle, T. L., and Putt, P. A.: Propylene and ethylene glycol as solvents for Sudan IV and Sudan Black B. Stain Tech., 26, (1951):51-56.
- Crossmon, G.: A modification of Mallory's connective tissue stain with a discussion of the principles involved. Anat. Rec., 69, (1937):33-38.
- Gershbein, L. L., and Spencer, K. L: Clinical studies in Aleutian disease of mink. Canadian J. Comp. Med. and Vet. Sci., 28, (1964):8-12.
- Gridley, M. F.: Manual of Histologic and Special Staining Techniques. Armed Forces Institute of Pathology, Washington, D. C., 1957.
- Hewer, T. F., Mathews, L. H., and Malkin, T.: Lipuria in tigers. Proc. Zool. Soc., London, 58, (1948): 326-328.

- Karstad, L.: Viral plasmacytosis (Aleutian disease) in mink: IV. Cytoplasmic glycoprotein inclusions and their differentiation from the viral inclusions of distemper." Can. J. Comp. Med. and Vet. Sci., 28, (1964):143-147.
- Karstad, L., and Pridham, T. J.: Aleutian disease of mink. I. Evidence of its viral etiology. Can.J. Comp. Med. and Vet. Sci., 26, (1962):97-102.
- Leader, R. W., Wagner, B. M., Henson, J., and Gorham, J.: Structural and histochemical observations of liver and kidney in Aleutian disease in mink. Amer. J. Path., 43, (1963):33-53.
- Leoschke, W. L.: Studies on the "Wet Belly" disease of mink. Proc. Ind. Acad. Sci., 72, (1962):136-318.
- Lillie, R. D.: Histologic Technique and Practical Histochemistry. McGraw-Hill Book Co., New York, 1954.
- Liu, H. C.: The comparative structure of the ureter. Amer. J. Anat., 111, (1962):1-15.
- Lobban, M. C.: Observations on intracellular lipid in the cat kidney. J. Anat., 89, (1955):92-99.
- Longley, J. B., Burtner, H. J., and Monis, B.: Mucous substances of excretory organs: A comparative study. Ann. N. Y. Acad. of Sci., 106, (1963):493-501.
- MacNider, W. B.: Occurrence of stainable lipoid material in renal epithelium of animals falling in different age groups. Proc. Soc. Exp. Biol. and Med., 58, (1945):326-328.
- Malewitz, T. D., and Smith, E. M: A nuclear stain employing dilute Harris hematoxylin. Stain Tech., 30,(1955): 311.
- Mallory, F. B.: Pathological Technique. W. B. Saunders Co., Philadelphia, 1944.
- Modell, W.: Observations on lipoids in the renal tubule of the cat. Anat. Rec., 57, (1933):13-27.
- Nielsen, I. M.: Urolithiasis in mink: pathology, bacteriology and experimental production. J. Urology, 75, (1956): 602.
- Schaible, P. J., Travis, H. F., and Shelts, G.: Urinary incontinence and "Wet Belly" in mink. Quar. Bull. of Mich. Ag. Exp. Sta., 44, (1962):466-483.

- Stranack, F.: Fat distribution in the kidney tubules of the Felidae. Proc. Zool. Soc. (London), 139, (1962): 475-482.
- Thompson, G. R., and Aliferis, P.: A clinical-pathological study of Aleutian mink disease: An experimental model for study of connective tissue disease. Arthritis Rheumatism, 7, (1964):521-533.
- Trautmann, A., and Fiebiger, J.: Fundamentals of the Histology of Domestic Animals. Translated and revised by Habel and Biberstein, Comstock Pub. Assoc., Ithaca, New York, 1957.
- Tucker, E., Lupton, C. H., and McManus, J. F. A.: A new inclusion in the visceral epithelium of the renal pelvis. Cancer, 12, (1959):1052-1059.
- Yadava, R. C. P., and Calhoun, M. L: Comparative histology of the kidney of domestic animals. Amer. J. Vet. Res., 19, (1958):958-968.

Figure 1.--Kidney Capsule

- 1. Outer adipose connective tissue portion of the kidney capsule.
- 2. Inner fibrous connective tissue portion of the kidney capsule.
- 3. Kidney cortex.



Figure 2.--Reticular network of the kidney and its capsule. Gridley's modified reticular stain x 200



Figure 3.--Kidney cortex

- 1. Macula densa
- 2. Juxtaglomerular apparatus

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- 3. Glomerulus
- 4. Proximal convoluted tubule
- 5. Distal convoluted tubule

Hematoxylin and eosin x 300

Figure 4.--Kidney cortex

- 1. Macula densa
- 2. Brush border of the proximal convoluted tubule

P.A.S. x 480





Figure 5.--Branching arcuate artery near the cortico-medullary junction of the kidney.

- 1. Arcuate artery
- 2. Arcuate vein
- 3. Proximal convoluted tubule with a prominent brush border
- P.A.S. x 200



Figure 6.--Concentrations of lipid material in the proximal convoluted tubules and thick descending limbs of Henle.

The dark rim around the tubules is composed of Oil Red O positive lipid material. See next figure for higher power.

Oil Red O x 32



- Figure 7.--Lipid droplets concentrated in the proximal convoluted tubules and thick descending limbs of Henle.
  - 1. Proximal convoluted tubule
  - 2. Thick descending limb of Henle
  - 3. Thin loop of Henle
  - 4. Thick ascending limb of Henle
  - Oil Red O x 500
- Figure 8.--P.A.S. positive material in the epithelium covering the renal crest.
  - 1. Epithelial covering the renal crest
  - 2. Papillary ducts
  - 3. Lumen of the renal pelvis
  - P.A.S. x 800



Figure 9.--P.A.S. positive intracytoplasmic granules in the epithelium lining the papillary ducts.

- 1. Papillary duct
- P.A.S. x 320

Figure 10.--P.A.S. positive intracytoplasmic granules in the epithelium lining the papillary ducts.

- 1. Papillary duct epithelium
- 2. P.A.S. positive granules

P.A.S. x 800



Figure 11.--Proximal ureter with surrounding fat tunic.

- 1. Inner longitudinal smooth muscle layer
- 2. Middle circular smooth muscle layer
- 3. Outer longitudinal smooth muscle layer
- 4. Surrounding fat tunic



Figure 12.--Dense reticular network in the lamina propria submucosa of the ureter.

Gridley's modified reticular stain x 300



Figure 13.--Muscular arrangement of the tunica muscularis in the proximal ureter.

- 1. Inner longitudinal muscle layer
- 2. Middle circular muscle layer
- 3. Outer longitudinal muscle layer
- 4. Transitional epithelium



Figure 14.--Cross section of the middle of the ureter showing the rich vascular plexus in the adventitia and the arrangement of the tunica muscularis.

- 1. Lumen of the ureter
- 2. Lamina propria-submucosa
- 3. Inner longitudinal smooth muscle layer
- 4. Middle circular smooth muscle layer
- 5. Outer longitudinal smooth muscle layer
- 6. Vascular plexus

Hematoxylin and eosin x 200



Figure 15.--Cross section of the distal ureter showing the marked thickening of the outer longitudinal muscle layer and the diminished inner longitudinal muscle layer

- 1. Inner longitudinal smooth muscle layer
- 2. Middle circular smooth muscle layer
- 3. Outer longitudinal smooth muscle layer



Figure 16.--Transitional epithelium lining the ureter

- 1. Transitional epithelium
- 2. Elongated spindle shaped cell with a hyperchromatic nucleus on the lumenal border
- 3. Capillary plexus just below the epithelium in the lamina propria-submucosa



Figure 17.--Transitional epithelium lining the ureter

1. Elongated spindle shaped cell on the lumenal surface of the ureter


Figure 18.--Dense reticular network of the bladder

- 1. Bladder lumen
- 2. Ureter

Gridley's modified reticular stain x 32

Figure 19.--Cross section of the bladder wall showing the arrangement of the tunica muscularis. In other areas the reverse arrangement is often true.

- 1. Inner longitudinal smooth muscle layer
- 2. Outer circular smooth muscle layer
- 3. Muscularis mucosae





Figure 20.--Urinary bladder mucosa

- 1. Transitional epithelium
- 2. Vascular plexus in the lamina propria
- 3. Capillary indenting the epithelium

Hematoxylin and eosin x 255

Figure 21.--Transitional epithelium of the bladder showing capillaries indenting its basal layers

1. Capillaries indenting the epithelium



Figure 22.--Bladder epithelium with surface cells about to be sloughed



Figure 23.--Cross section of the outer bladder wall

- 1. Adventitia (serosa)
- 2. Tunica muscularis
- 3. Autonomic ganglion

Hematoxylin and eosin x 75

Figure 24.--Cross Section of the prostatic urethra

- 1. Lumen of the prostatic urethrae
- 2. Ductus deferens



Figure 25.--Prostatic duct opening into the urethral lumen

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- 1. Prostatic duct
- 2. Urethral lumen



Figure 26.--Cavernous spaces in the lamina propria of the prostatic urethra

1. Cavernous spaces

Hematoxylin and eosin x 200

Figure 27.--Elastic fibers in the lamina propria of the prostatic urethra

1. Elastic fibers

Weigert VanGieson x 800



Figure 28.--Prostatic utriculus opening on the colliculus seminalis of the prostatic urethra

1. Prostatic utriculus

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2. Colliculus seminalis

Hematoxylin and eosin  $\mathbf{x}$  200



Figure 29.--Cross section of the membranous urethra

- 1. Urethral lumen
- 2. Cavernous spaces
- 3. Skeletal urethral muscle

Hematoxylin and  $eosin \times 30$ 

Figure 30.--Mucosa of the membranous urethra

1. Urethral lumen

Hematoxylin and eosin x 120

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Figure 31.--Epithelial lining of the membranous urethra Hematoxylin and eosin x 200



Figure 32.--Cross section of the membranous urethra showing the extensive network of elastic fibers in the lamina propria

1. Lamina propria

Weigert VanGieson x 30

Figure 33.--Cross section of the membranous urethra showing the extensive network of elastic fibers in the lamina propria

- 1. Lamina propria
- 2. Elastic fibers

Weigert VanGieson x 480





Figure 34.--Cross section of the proximal penile urethral mucosa

Numerous venous sinuses in the lamina propria

Hematoxylin and eosin x = 200

Figure 35.--Epithelial lining of the proximal penile urethra

- 1. Stratified columnar epithelium
- 2. Intraepithelial nest of mucous cells
- 3. Lacunae of Morgagni





Figure 36.--Cross section of the penile urethra near the external urethral orifice

- 1. Os penis
- 2. Urethral lumen
- 3. Venous sinuses engorged with blood
- 4. Nerve plexus

Hematoxylin and  $eosin \times 30$ 

Figure 37.--Epithelial lining of the penile urethra near the external urethral orifice



Figure 38.--Cross section of the female urethra showing the dense reticular network in the lamina propria-submucosa

1. Reticular fibers

Gridley's modified reticular stain x 30

Figure 39.--Cross section of the proximal female urethra

- 1. Skeletal urethral muscle
- 2. Urethral lumen
- 3. Lacunae of Morgagni

Hematoxylin and  $eosin \times 30$ 



Figure 40.--Female urethral epithelium (mid portion)

- 1. Lacunae of Morgagni
- 2. Intraepithelial nest of mucous glands

Hematoxylin and eosin x 120

Figure 41.--Nests of serous glands located deep in the submucosa and between the bands of the skeletal urethral muscle of the female urethra

- 1. Nests of serous glands
- 2. Skeletal urethral muscle
- 3. Submucosa



Figure 42.--Intraepithelial cysts in the female urethral epithelium

- 1. Intraepithelial cyst
- 2. Lacunae of Morgagni
- 3. Nests of mucous cells

Hematoxylin and eosin x 120

Figure 43.--Stratified sqamous epithelium lining the distal female urethra

Hematoxylin and  $eosin \times 300$ 

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