

CMHD Pathology Core Toronto Centre for Phenogenomics 25 Orde St. 3rd fl. Toronto, Ont. M5T 3H7 Tel.(416) 586-8375 Fax (416) 586-5993

contact: Dr. Susan Newbigging email: newbigging@lunenfeld.ca

CMHD Pathology Report

Principle Investigator: Dr. Jacqui White Institute: Wellcome Trust Sanger Institute Address: Attn: Linda Read Wellcome Trust Genome Campus Hinxton Cambridge CB10 1SA, UK

ReportID: Report Date: June 20, 2013 Pathologist: Dr. H. Adissu



Mouse Genetics Project

Wellcome Trust Sanger Institute Wellcome Trust Genome Campus Hinxton, Cambridge CB10 1SA UK email: MGPenquiries@sanger.ac.uk Mouse Portal Europhenome

CMHD LabID: N13-469

Relevant History:

Phenotype abnormal fertility/fecundity

AnimalID: M00351452 (Male) **Histopathology Findings:**

testis (MA:0000411)

Histopath Description:

Numerous seminiferous tubules are vacuolated. Nearly all seminiferous tubules are devoid of maturing spermatids and spermatocytes. The epididymis contains abundant cellular debri and proteinaceous fluid with rare spermatocytes.

Morphological Diagnosis:

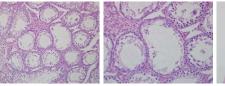
Distribution: generalized; Severity: severe; MPATH Diagnosis: degenerative change MPATH:14

Definitive Diagnosis:

Seminiferous tubule, vacuolation and atrophy - absence of spematocytogensis

Histopathology Comments:

The lesion is consistent with abnormal fertility.



Seminiferous tubule, Seminiferous tubule, Epididymal duct, vacuolation and vacuolation and atrophy - absence atrophy - absence of of spematocytogensis, spematocytogensis, 20x.HF



40x, HE

20x, HE

liver (MA:0000358)

Histopath Description: diffuse lipidosis

Morphological Diagnosis: Distribution: diffuse; Severity: severe; MPATH Diagnosis: steatosis MPATH:622

Definitive Diagnosis: Hepatic lipidosis

brain (MA:0000168) **Histopath Description:** There is mild dilation of the lateral ventricles Morphological Diagnosis:

Distribution: bilateral; Severity: mild;

www.cmhd.ca/pathology/reports/histopathology_report_wtsi.asp?ID=83680855

Definitive Diagnosis: Dilation of the brain ventricles

Histopathology Comments:

Mild dilation of the lateral ventricles is a background condition in mice of C57BL/6N background (Brayton et al., 2004).

stomach (MA:0000353)

Histopath Description:

mild neutrophilic gastritis

Morphological Diagnosis:

Distribution: multifocal to coalescing; Severity: mild;

Definitive Diagnosis:

Gastrits, neutrophilic

Organ/Tissue Analyzed:

Histopathology examination included the following organs and tissues: brain, trigeminal ganglion, eyes, salivary glands, trachea, lungs, heart, thymus, thyroid gland, parathyroid gland, exocrine and endocrine pancreas, oesophagus, stomach, small intestine, large intestine, liver, gall bladder, spleen, kidneys, adrenal gland, lymph nodes, spinal cord, bone marrow, sternum, femur and tibia with associated skeletal muscles, brown fat, pinna, skin, testis, epididymis, seminal vesicle, and prostate.

AnimalID: M00351454 (Male)

Histopathology Findings:

testis (MA:0000411)

Histopath Description:

Numerous seminiferous tubules are vacuolated. Nearly all seminiferous tubules are devoid of maturing spermatids and spermatocytes. The epididymis contains abundant cellular debri and proteinaceous fluid with rare spermatocytes.

Morphological Diagnosis:

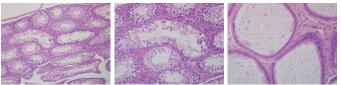
Distribution: generalized; **Severity:** severe; **MPATH Diagnosis:** degenerative change MPATH:14

Definitive Diagnosis:

Seminiferous tubule, vacuolation and atrophy - absence of spematocytogensis

Histopathology Comments:

The lesion is consistent with abnormal fertility.



Seminiferous tubule, Seminiferous tubule, Epididymal duct,
vacuolation and
atrophy - absence
ofepididymal
aspermia/hypospermia,
40x, HEspematocytogensis,
20x, HE40x, HE

liver (MA:0000358)

Histopath Description: diffuse lipidosis

Morphological Diagnosis:

Distribution: diffuse; Severity: severe; MPATH Diagnosis: steatosis MPATH:622 Definitive Diagnosis:

Hepatic lipidosis

brain (MA:0000168)

Histopath Description: There is mild dilation of the lateral ventricles

Morphological Diagnosis: Distribution: bilateral; Severity: mild;

Definitive Diagnosis: Dilation of the brain ventricles

Histopathology Comments: Mild dilation of the lateral ventricles is a background condition in mice of C57BL/6N background (Brayton et al., 2004).

stomach (MA:0000353)

Histopath Description:

mild neutrophilic gastritis

Morphological Diagnosis:

Distribution: multifocal to coalescing; Severity: mild;

Definitive Diagnosis:

Gastrits, neutrophilic

retina (MA:0000276)

```
Histopath Description:
```

Involving one eye, there are clusters of external nuclear structures within the internal plexiform layer.

Morphological Diagnosis:

Distribution: Focal; Severity: mild;

Definitive Diagnosis: Retinal dysplasia

Organ/Tissue Analyzed:

Histopathology examination included the following organs and tissues: brain, trigeminal ganglion, eyes, salivary glands, trachea, lungs, heart, thymus, thyroid gland, parathyroid gland, exocrine and endocrine pancreas, oesophagus, stomach, small intestine, large intestine, liver, gall bladder, spleen, kidneys, adrenal gland, lymph nodes, spinal cord, bone marrow, sternum, femur and tibia with associated skeletal muscles, brown fat, pinna, skin, testis, epididymis, seminal vesicle, and prostate.

AnimalID: M00236621 (Female) **Histopathology Findings:** liver (MA:0000358) **Histopath Description:** diffuse lipidosis Morphological Diagnosis: Distribution: diffuse; Severity: severe; MPATH Diagnosis: steatosis MPATH:622 **Definitive Diagnosis:** Hepatic lipidosis parathyroid gland (MA:0000128) **Histopath Description:** There are large well demarcated clusters of lymphoid cells within the parathyroid gland Morphological Diagnosis: Distribution: multifocal; **Definitive Diagnosis:** Ectopic thymic tissue **Histopathology Comments:** incidental

brain (MA:0000168) Histopath Description: There is mild dilation of the lateral ventricles

Morphological Diagnosis: Distribution: bilateral; Severity: mild; Definitive Diagnosis: Dilation of the brain ventricles Histopathology Comments: Mild dilation of the lateral ventricles is a background condition in mice of C57BL/6N background (Brayton et al., 2004).

Organ/Tissue Analyzed:

Histopathology examination included the following organs and tissues: brain, trigeminal ganglion, eyes, salivary glands, trachea, lungs, heart, thymus, thyroid gland, parathyroid gland, exocrine and endocrine pancreas, oesophagus, stomach, small intestine, large intestine, liver, gall bladder, spleen, kidneys, adrenal gland, lymph nodes, spinal cord, bone marrow, sternum, femur and tibia with associated skeletal muscles, brown fat, pinna, skin, uterus, oviduct, and ovary, and mammary gland.

AnimalID: M00236623 (Female) Histopathology Findings: liver (MA:0000358)

Histopath Description:

minimal lipidosis Morphological Diagnosis:

Distribution: multifocal; Severity: mild;

Definitive Diagnosis:

Hepatic lipidosis, minimal

lymph node (MA:0000139)

Histopath Description:

The mesenteric lymph node is enlarged (greater than three-fold). There are multiple follicles with large germinal centers. The sinuses contain large numbers of mature lymphocytes.

Morphological Diagnosis:

Duration: Sub-acute; **Distribution:** Diffuse; **Severity:** moderate; **MPATH Diagnosis:** hyperplasia MPATH:134

Definitive Diagnosis:

Lymphoid hyperplasia.

Histopathology Comments:

The changes in the mesenteric lymph node are suggestive of draining of a regional inflammatory process. However, such a process was not observed in the tissues examined.

brain (MA:0000168)

Histopath Description: There is mild dilation of the lateral ventricles. The periventricular neurpil is rarefied

Morphological Diagnosis:

Distribution: bilateral; Severity: mild;

Definitive Diagnosis:

Dilation of the brain ventricles with rarefaction of the periventricular neuropil

Histopathology Comments:

Mild dilation of the lateral ventricles is a background condition in mice of C57BL/6N background (Brayton et al., 2004).

spleen (MA:0000141)

Histopath Description:

marked erythroid hyperplasia

Morphological Diagnosis:

Distribution: multifocal; **Severity:** moderate; **MPATH Diagnosis:** extramedullary hemopoiesis MPATH:595

Definitive Diagnosis:

Splenic erythroid hyperplasia

Organ/Tissue Analyzed:

Histopathology examination included the following organs and tissues: brain, trigeminal ganglion, eyes, salivary glands, trachea, lungs, heart, thymus, thyroid gland, parathyroid gland, exocrine and endocrine pancreas, oesophagus, stomach, small intestine, large intestine, liver, gall bladder, spleen, kidneys, adrenal gland, lymph nodes, spinal cord, bone marrow, sternum, femur and tibia with associated skeletal

muscles, brown fat, pinna, skin, uterus, oviduct, and ovary, and mammary gland.

Report Summary and Recommendation:

Testicular atrophy with lack of spermatocytogenesis is evident in both males consistent with infertility documented in this line. Other lesions are attributable to diet or strain background.

Line summary: Testicular degeneration and atrophy; epidydimal hypospermia (2/2)