

CMHD Pathology Report



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

Pathologist: H. Adissu

ReportID: Report Date: November 23,

2011

Mouse Genetics Project Wellcome Trust Sanger

Institute Wellcome Trust Genome Campus Hinxton, Cambridge CB10 1SA UK

CMHD LabID: N11-85

Relevant History:

(Viability at postnatal day 14, Recessive Lethal Study (E14.5)) Embryonic lethality, pre-weaning lethality

AnimalID: M00431715-F730047E07Rik Het

Tissue Preservation and Staining:

Tissues not present in submission: Calvarium, ears, tongue, Harderian gland, zymbal gland, nasal sinuses, teeth, gall bladder.

Histopathology Findings:

eye (MA:0000261)

Histopath Description:

A transverse section of the eye at the anterior portion is present (evidenced by circumferential presence of the lenticular epithelium). Only a small segment of the retina (0.5 mm in length) is present in this section. This segment of the retina has thin layers of external and inner nuclear layers (nearly half normal thickness). The outer plexiform layer is attenuated causing multifocal blending of the nuclear layers. There is a focally extensive retinal detachment at one edge of this retinal segment; few retinal epithelial cells are hypertrophic and cuboidal (tombstone appearance). Ganglion cells are rare within this segment.

Morphological Diagnosis:

Distribution: Unilateral; **Severity:** moderate;

Definitive Diagnosis:

Segmental retinal dysplasi with focally xtensive retinal detachment.

Histopathology Comments:

This section was not optimal for complete evaluation of all ocular structures.



Eye, transverse anterior section 10x, HE.

stomach (MA:0000353)

Histopath Description:

There are low numbers of neutrophils within the deep lamina propria

Morphological Diagnosis:

Duration: Sub-acute; Distribution: Multifocal; Severity: moderate; MPATH Diagnosis: inflammation MPATH:212

Definitive Diagnosis:

Gastritis, suppurative

Histopathology Comments:

This lesion is most commonly associated with Helicobacter infection. Further investigation is suggested using histochemistry (Silver stain) or colony fecal PCR.

liver (MA:0000358)

Histopath Description:

The overall hepatic lobular architecture is normal. Diffusely, hepatocytes contain intracytoplasmic clear vacuoles (lipid). The lipid vacuoles within the midzonal and periacinar regions are small (2-3 um in diameter) and surround a central nucleus (interpreted as microvesicular lipid). The lipid vacuoles within the portal areas are large (8-12 um in diameter) and displace the nucleus to the margin (macrovesicular lipid).

Morphological Diagnosis:

Distribution: Diffuse; Severity: moderate; MPATH Diagnosis: lipid deposition MPATH:42

Definitive Diagnosis:

Hepatic lipidosis

Histopathology Comments:

Hepatocellular vacuolar change of variable degree suggestive of lipidosis is present in all mice from WTSI, consistent with high lipid diet.

lymph node (MA:0000139)

Histopath Description:

The mesenteric lymph node is markedly enlarged. Lymphoid follicles are enlarged and there are numerous germinal centers. Numerous mature lymphocytes are present within the medullary and paracortical sinuses. The marginal sinuses contain numerous mature lymphocytes.

Morphological Diagnosis:

Severity: moderate; MPATH Diagnosis: hyperplasia MPATH:134

Definitive Diagnosis:

Lymphoid hyperplasia

salivary gland (MA:0000346)

Histopath Description:

Within the mandibular salivary gland, the interstitium (mainly surrounding blood vessels) are expanded by multifocal aggregates of lymphocytes, histiocytes, and low numbers of plasma cells.

Morphological Diagnosis:

Duration: Chronic-active; **Distribution:** Multifocal; **Severity:** mild; **MPATH Diagnosis:** inflammation MPATH:212

Definitive Diagnosis:

Interstitial histiocytic and lymphocytic sialadenitis

Histopathology Comments:

This is a common and insignificant incidental finding in mice.

brain (MA:0000168)

Histopath Description:

There is a mild enlargement of the third ventricle.

Morphological Diagnosis:

Severity: mild; **MPATH Diagnosis:** hydrocephalus MPATH:639

Definitive Diagnosis:

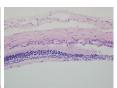
hydrocephalus, third ventricle

Histopathology Comments:

Variable degree of hydrocephalus is observed in a proportion of wild type C57 Black 6 mice.







Retina, transverse Retina, detachment Retina, detachment anterior section, 4x, and dysplasia, 20x, and dysplasia, 40x, HE. HE

Organ/Tissue Analyzed:

There were no significant findings in the following tissues: Trachea, lungs, heart, thymus, thyroid gland, parathyroid gland, spleen, exocrine and endocrine pancreas, esophagus, stomach, intestines, urinary organs and tract, adrenal gland, reproductive organs, spinal cord, bones, bone marrow, skeletal muscles, brown fat, and skin.

AnimalID: M00431716-F730047E07Rik Het

Tissue Preservation and Staining:

Tissues not present in submission: Calvarium, ears, tongue, Harderian gland, zymbal gland, nasal sinuses, teeth, gall bladder.

Histopathology Findings:

brain (MA:0000168)

Histopath Description:

There is a mild enlargement of the third ventricle.

Morphological Diagnosis:

Severity: mild; MPATH Diagnosis: hydrocephalus MPATH:639

Definitive Diagnosis:

hydrocephalus, third ventricle

Histopathology Comments:

Variable degree of hydrocephalus is observed in a proportion of wild type C57 Black 6 mice.

stomach (MA:0000353)

Histopath Description:

There are low numbers of neutrophils within the deep lamina propria. There are rare ectatic glands containing cell debri.

Morphological Diagnosis:

Duration: Sub-acute; **Distribution:** Multifocal; **Severity:** moderate; **MPATH Diagnosis:** inflammation MPATH:212

Definitive Diagnosis:

Gastritis, suppurative

Histopathology Comments:

This lesion is most commonly associated with Helicobacter infection. Further investigation is suggested using histochemistry (Silver stain) or colony fecal PCR.

liver (MA:0000358)

Histopath Description:

The overall hepatic lobular architecture is normal. Diffusely, hepatocytes contain intracytoplasmic clear vacuoles (lipid). The lipid vacuoles within the midzonal and periacinar regions are small (2-3 um in diameter) and surround a central nucleus (interpreted as microvesicular lipid). The lipid vacuoles within the portal areas are large (8-12 um in diameter) and displace the nucleus to the margin (macrovesicular lipid).

Morphological Diagnosis:

Distribution: Diffuse; Severity: moderate; MPATH Diagnosis: lipid deposition MPATH:42

Definitive Diagnosis:

Hepatic lipidosis

Histopathology Comments:

Hepatocellular vacuolar change of variable degree suggestive of lipidosis is present in all mice from WTSI, consistent with high lipid diet.

lung (MA:0000415)

Histopath Description:

There is a focal perivascular mononuclear inflammatory cell aggregates within the lung

Morphological Diagnosis:

Duration: Chronic; **Distribution:** Focal; **Severity:** mild; **MPATH Diagnosis:** inflammation MPATH:212

Definitive Diagnosis:

Perivascular inflammatory aggregates

Histopathology Comments:

This lesion is suggestive of antigenic stimulation of hematogenous origin. It is a common and insignificant incidental finding.



Eye, normal, 4x, HE

Organ/Tissue Analyzed:

There were no significant findings in the following tissues: Eyes, salivary glands, trachea, heart, thymus, thyroid gland, parathyroid gland, spleen, exocrine and endocrine pancreas, esophagus, intestines, urinary organs and tract, adrenal gland, reproductive organs, lymph nodes, spinal cord, bones, bone marrow, skeletal muscles, brown fat, and skin.

AnimalID: M00370329-F730047E07Rik Het

Tissue Preservation and Staining:

Tissues not present in submission: Calvarium, ears, tongue, Harderian gland, zymbal gland, nasal sinuses, teeth, gall bladder. One of the eyes is obliquely sectioned.

Histopathology Findings:

brain (MA:0000168)

Histopath Description:

There is a mild enlargement of the third ventricle.

Morphological Diagnosis:

Severity: mild; **MPATH Diagnosis:** hydrocephalus MPATH:639

Definitive Diagnosis:

hydrocephalus, third ventricle

Histopathology Comments:

Variable degree of hydrocephalus is observed in a proportion of wild type C57 Black 6 mice.

liver (MA:0000358)

Histopath Description:

The overall hepatic lobular architecture is normal. Diffusely, hepatocytes contain intracytoplasmic clear vacuoles (lipid). The lipid vacuoles within the midzonal and periacinar regions are small (2-3 um in diameter) and surround a central nucleus (interpreted as microvesicular lipid). The lipid vacuoles within the portal areas are large (8-12 um in diameter) and displace the nucleus to the margin (macrovesicular lipid).

Morphological Diagnosis:

Distribution: Diffuse; Severity: moderate; MPATH Diagnosis: lipid deposition MPATH:42

Definitive Diagnosis:

Hepatic lipidosis

Histopathology Comments:

Hepatocellular vacuolar change of variable degree suggestive of lipidosis is present in all mice from WTSI, consistent with high lipid diet.



Eye, normal, optic nerve not in view, 4x, HE.

Organ/Tissue Analyzed:

There were no significant findings in the following tissues: Eyes, salivary glands, trachea, lungs, heart, thymus, thyroid gland, parathyroid gland, spleen, exocrine and endocrine pancreas, esophagus, stomach, intestines, urinary organs and tract, adrenal gland, reproductive organs, lymph nodes, spinal cord, bones, bone marrow, skeletal muscles, brown fat, and skin.

AnimalID: M00360999-F730047E07Rik Het

Tissue Preservation and Staining:

Tissues not present in submission: Calvarium, ears, tongue, Harderian gland, zymbal gland, nasal sinuses, teeth, gall bladder. Only a section of the cornea is present for one of the eyes.

Histopathology Findings:

eye (MA:0000261)

Histopath Description:

Only one of the eyes is examined (the other eye is at a deep level in the block). Clusters of nuclei within the outer plexiform layer; the nuclei resemble that of of rods and cons.

Morphological Diagnosis:

Distribution: Unilateral; **Severity:** mild; **MPATH Diagnosis:** developmental and structural abnormality MPATH:55

Definitive Diagnosis:

Retinal dysplasia - ectopic rods and cons

brown fat (MA:0000057)

Histopath Description:

Within the cervical brown adipose tissue, there are small aggregates of inflammatory cells composed of monocytes, lymphocytes, and rare neutrophils. The lipocytes in the vicinity have a central prominent round open nucleus and the cytoplasm contain numerous small lipid vacuoles.

Morphological Diagnosis:

Duration: Chronic-active; **Distribution:** Multifocal; **Severity:** mild; **MPATH Diagnosis:** inflammation MPATH:212

Definitive Diagnosis:

Steatitis, lymphocytic and histiocytic.

liver (MA:0000358)

Histopath Description:

The overall hepatic lobular architecture is normal. Diffusely, hepatocytes contain intracytoplasmic clear vacuoles (lipid). The lipid vacuoles within the midzonal and periacinar regions are small (2-3 um in diameter) and surround a central nucleus (interpreted as microvesicular lipid). The lipid vacuoles within the portal areas are large (8-12 um in diameter) and displace the nucleus to the margin (macrovesicular lipid).

Morphological Diagnosis:

Distribution: Diffuse; Severity: moderate; MPATH Diagnosis: lipid deposition MPATH:42

Definitive Diagnosis:

Hepatic lipidosis

Histopathology Comments:

Hepatocellular vacuolar change of variable degree suggestive of lipidosis is present in all mice from WTSI, consistent with high lipid diet.

salivary gland (MA:0000346)

Histopath Description:

The interstitium (mainly surrounding blood vessels) is multifocally expanded by agreggates of lymphocytes, histiocytes, and low numbers of plasma cells.

Morphological Diagnosis:

Duration: Chronic-active; **Distribution:** Multifocal; **Severity:** mild; **MPATH Diagnosis:** inflammation MPATH:212

Definitive Diagnosis:

Interstitial histiocytic and lymphocytic sialadenitis

Histopathology Comments:

This is a common and insignificant incidental finding in mice.

skin (MA:0000151)

Histopath Description:

There is a focal two-fold epidermal hyperplasia. There are low numbers of perivascular mononuclear inflammatory cells in the underlying dermis.

Morphological Diagnosis:

Duration: Chronic; **Distribution:** Focal; **Severity:** mild; **MPATH Diagnosis:** hyperplasia MPATH:134

Definitive Diagnosis:

Epidermal hyperplasia

lung (MA:0000415)

Histopath Description:

There is a prominent focus of perivascular mononuclear inflammatory cell aggregate within the lung **Morphological Diagnosis:**

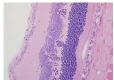
Duration: Chronic; **Distribution:** Focal; **Severity:** mild; **MPATH Diagnosis:** inflammation MPATH:212

Definitive Diagnosis:

Perivascular inflammatory aggregates

Histopathology Comments:

This lesion is suggestive of antigenic stimulation of hematogenous origin. It is a common and insignificant incidental finding.



Retina, dysplasia, note clusters of nuclei of rods and cons in outer plexiform layer, 40x, HE.

Fat, steatitis, 40x,

Organ/Tissue Analyzed:

There were no significant findings in the following tissues: Brain, trachea, heart, thymus, thyroid gland, parathyroid gland, spleen, liver, exocrine and endocrine pancreas, esophagus, stomach, intestines, urinary organs and tract, adrenal gland, reproductive organs, lymph nodes, spinal cord, bones, bone marrow, and skeletal muscles.

Summary:

Retinal abnormality is observed in two mice (M00360999 and M00431715).

The dysplastic changes in M00360999 is mild. A more severe of version of this lesion was noted in one of the wild type controls (M00405977-Snip1WT), suggesting that it may be a strain background lesion

Report Summary and Recommendation:

Retinal abnormality is observed in two mice (M00360999 and M00431715).

The dysplastic changes in M00360999 is mild. A more severe of version of this lesion was noted in one of the wild type controls (M00405977-Snip1WT), suggesting that it may be a strain background lesion. Recently a mutation in Crb1 is confirmed rd8 mutation in the Crb1 gene in C57BL/6N substrain is associated with multiple light-colored spots in the fundus of the eye that correspond histologically to retinal folds, pseudorosettes, and focal retinal dysplasia and degeneration. Hence retinal lesions in this line may be a background lesion.

Retina: dysplasia MPATH:64