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CTE CENTER NEUROPATHOLOGY REPORT

K0623

PATIENT'S NAME: Ellison, Kevin
AUTOPSY#: K-0623
DATE OF DEATH: 10/04/2018
DATE OF AUTOPSY: Unknown
DATE BRAIN RECEIVED: 1/22/2019
FROM: Los Angeles, CA
TYPE OF SPECIMEN: Fresh whole brain, dura, pituitary
Brain weight: 1374g

FINAL DIAGNOSES:

1. Chronic Traumatic Encephalopathy (CTE): Stage II

Comment: There is mild ventricular dilation and frontal lobe atrophy. There is a small cavum septum pellucidum. Multiple perivascular CTE lesions are present in the superior frontal, dorsolateral frontal, inferior temporal and inferior parietal cortices. Superficial neurofibrillary tangles (NFTs) are also found in the temporal cortices. There is mild neurofibrillary degeneration of the locus coeruleus and substantia nigra. These changes conform to Chronic Traumatic Encephalopathy (CTE), Stage II (out of a possible IV, with IV being the most severe).

McKee AC, Cantu RC, Nowinski CJ, Hedley-Whyte ET, Gavett BE, Budson AE, Santini VE, Lee H-Y, Kubilus CA, Stern RA. Chronic Traumatic Encephalopathy in Athletes: Progressive Tauopathy following Repetitive Head Injury. *J Neuropath Exp Neurol*, 2009 68(7): 709-735.

McKee AC, Stein TD, Nowinski CJ, Stern RA, Daneshvar DH, Alvarez VE, Lee H-S, Hall GF, Wojtowicz SM, Baugh CM, David O. Riley DO, Kubilus CA, Cormier KA, Jacobs MA, Martin BR, Abraham CR, Ikezu T, Reichard RR, Wolozin BL, Budson AE, Goldstein LE, Kowall NW, Cantu RC. The Spectrum of Disease in Chronic Traumatic Encephalopathy, Brain 2013, Jan; 136(Pt 1): 43-64. doi: 10.1093/brain/aws307.

McKee AC, Cairns NJ, Dickson DW, Folkerth RD, Keene CD, Litvan I, Perl DP, Stein TD, Stewart W, Vonsattel JP, Tripodis Y, Alvarez VE, Bieniek KF, Crary J, Dams-O'Connor K, Gordon W and the TBI/CTE group. The First NINDS/NIBIB Consensus Meeting to Define Neuropathological Criteria for the Diagnosis of Chronic Traumatic Encephalopathy. *Acta Neuropathol*. 2016 Jan; 131(1): 75-86. doi: 10.1007/s00401-015-1515-z.

Mez J, Daneshvar D, Kiernan P, Abdolmohammadi B, Alvarez V, Huber B, Alosco M, Solomon T, Nowinski C, McHale L, Cormier K, Kubilus C, Martin B, Murphy L, Baugh C, Montenegro P, Chaisson C, Tripodis Y, Kowall N, Weuve J, McClean M, Cantu R, Goldstein L, Katz D, Stern R, Stein T, **McKee A**. Clinicopathological Evaluation of Chronic Traumatic Encephalopathy in Players of American Football. *JAMA*. 2017 Jul 25; 318(4): 360-370. doi: 10.1001/jama.2017.8334. PMID: 28742910

Montine TJ, Phelps CH, Beach TG, Bigio EH, Cairns NJ, Dickson DW, Duyckaerts C, Frosch MP, Masliah E, Mirra SS, Nelson PT, Schneider JA, Thal DR, Trojanowski JQ, Vinters HV, Hyman BT. National Institute on Aging-Alzheimer's Association guidelines for the neuropathologic assessment of Alzheimer's disease: a practical approach. *Acta Neuropathol* 2012; 123:1-11. PMC3268003

GROSS EXAMINATION

The meninges are translucent

The cranial nerves (I-XII): unremarkable

The cerebral blood vessels: unremarkable

Atherosclerosis of cerebral vessels: none

There are no contusions.

There is no atrophy of the cerebral cortex, basal ganglia, brainstem or cerebellum

Cavum septum: 1.0 cm AP x 0.3 cm lateral x 0.8 cm DV

Ventricular size:

Lateral ventricles: 1+ enlarged

Third ventricle: 0-1+ enlarged

Fourth ventricle: unremarkable

Cerebral aqueduct: unremarkable

Parenchymal vascular lesions:

Infarct(s) (>1.0 cm in diameter): none

Lacunes (<1.0 cm): none

Hemorrhages (parenchymal): none

There is no atrophy of hippocampal formation, amygdala, entorhinal cortex, striatum, hypothalamus, mammillary bodies, or thalamus.

Brainstem:

Substantia nigra: unremarkable

Locus coeruleus: 1+ pallor

Cerebellum: unremarkable

Other non-vascular gross CNS: none

MICROSCOPIC EXAMINATION

Available for microscopic examination are sections from representative regions listed below. A subset of slides was prepared from the paraffin embedded blocks and stained with Luxol fast blue, hematoxylin and eosin (LHE), and some of the following: AT8, Alpha-synuclein, Amyloid beta, TDP-43, PrP and APP

Key sheet to possible sections

1. Olfactory bulb
2. Midbrain at level of red nucleus
- 2A. Midbrain at superior cerebellar peduncle
3. Precentral, postcentral cortex (BA 4,3,2,1)
4. Inferior parietal cortex (BA 39,40)
5. Anterior cingulate (BA 24)
- 5A. Superior frontal (BA 8,9)
6. Inferior frontal cortex (BA 10,11,12)
7. Dorosolateral frontal (BA45, 46)
- 8A. Caudate, putamen, and accumbens (CAP) with septal cortex, fornix
- 8B. Insular cortex
9. Anterior temporal (BA 38)
10. Superior temporal (BA 20, 21,22)
11. Amygdala, with entorhinal cortex (BA 28)
12. Globus pallidus, insula, sub. Innominata
13. Anterior hippocampus
14. Hippocampal formation, lateral geniculate
15. Superior temporal posterior (BA 41,42)

16. Thalamus with subthalamic nucleus
- 16A Thalamus with mammillary body
17. Posterior cingulate (BA23, 31)
18. Calcarine cortex (BA 17,18)
19. Superior parietal cortex (BA 7B)
20. Upper pons (level of locus coeruleus)
- 20A. Lower pons at Vth cranial nerve
21. Medulla oblongata (including inferior olives)
- 22-1. Cervical spinal cord
- 22-2,3 Thoracic spinal cord
- 22-4, 5 Lumbar spinal cord
- 22-6 Sacral spinal cord
23. Cerebellar vermis
24. Cerebellum with dentate nucleus
25. Parastriate cortex

MICROSCOPIC FINDINGS

I. Leptomeninges:

Fibrosis: none

III. Cerebral Blood Vessels:

Arteriolosclerosis: mild

Amyloid angiopathy:

Leptomeninges: none

Intraparenchymal: none

IV. Cerebral cortex:

Cytoarchitecture (radial and laminar): normal

Neuronal loss: none

Spongiform change: none

NFTs: (AT8)

Rolandic: perivascular lesion at depth of sulcus

Superior frontal: perivascular lesions at depth of sulcus

Dorsolateral frontal: perivascular lesions at depth of sulcus

Inferior temporal: perivascular lesions at depth of sulcus

Inferior parietal: perivascular lesions at depth of sulcus

Temporal pole: superficial NFTs

Temporal isocortex: superficial NFTs

Calcarine: none

A β /Bielschowsky

SPs: (diffuse): none

SPs: (neuritic): none

TDP-43: Dorsolateral frontal: none

PrP: none

Hippocampal formation: unremarkable

Entorhinal cortex: unremarkable

Cerebral white matter:

Loss of myelinated nerve fibers: mild

Arteriolosclerosis: mild

Microinfarcts: none

Perivascular macrophages: 2-3+

V. Subcortical Nuclei:

Amygdala:

NFTs: 2+

Thalamus: 1+ NFTS

VI. Brainstem

Substantia nigra, pars compacta:

Neuronal loss: none

Lewy bodies: none

NFTs: 1+

TDP-43: none

Locus coeruleus:

NFTs: 1+

VII. Cerebellum:

Cortex: unremarkable

Dentate nucleus: unremarkable

Purkinje cells: unremarkable

White matter:

Astrocytosis: 2+

Myelin loss: 2+

VII: Spinal cord: unremarkable

NEUROPATHOLOGIST:

A handwritten signature in black ink, appearing to read "Ann C. McKee", written in a cursive style.

Ann C. McKee, MD