

Suprasellar germ cell tumor in a dog

Tumor supra-selar de células germinativas em um cão

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- NOTE -

ABSTRACT

A case of suprasellar germ cell tumor is described in a 6-year-old Airedale Terrier bitch that presented behavioral changes and abrupt onset of blindness. The neoplasm compressed the ventral surface of the cerebrum from the level of basal ganglia to the mesencephalon. Histologically the neoplasm consisted of nests and trabeculae of round to polygonal cells that occasionally surround tubules and cysts formed by columnar cells. Neoplastic cells are immunopositive for cytokeratin and alpha-fetoprotein. The diagnosis was based on location, histological appearance and on the immunohistochemical staining.

Key words: *extragonadal germ cell tumor, canine neoplasms, central nervous system.*

RESUMO

É descrito um caso de tumor de células germinativas supra-selar em uma cadela Airedale Terrier de seis anos de idade, que apresentou transtornos do comportamento e aparecimento abrupto de cegueira. O neoplasma comprimiu a superfície ventral do cérebro desde a altura dos núcleos basais até o mesencéfalo. Histologicamente, o neoplasma consistia de ninhos e trabéculas de células redondas ou poligonais que ocasionalmente arranjavam-se ao redor de túbulos e cistos formados por células colunares. As células neoplásicas foram positivas na imunoistoquímica para citoqueratina e α -fetoproteína. O diagnóstico foi feito com base na localização do tumor, no seu aspecto histológico e nos resultados da marcação imunoistoquímica.

Palavras-chave: *tumor de células germinativas extragonadal, neoplasmas de cães, sistema nervoso central.*

Extragenadal germ cell tumors can originate from the transformation of the otherwise normal resident population of germ cells, of a developmentally derived ectopic rest of germ cells, or of germ cells that migrated into the central nervous system late in development (FERREIRA et al., 2003; NYSKA et al., 1993). Usually, germ cells migrate from the wall of the yolk sac to the primitive gonad. As a rule, these germ cells do not survive and are eliminated by immune mechanisms; those cells which do survive may undergo neoplastic transformation (FERREIRA et al., 2003). Most extragenadal germ cell tumors in humans arise intracranially in midline diencephalic structures, i.e., the pineal region or the suprasellar-hypothalamo-hypophyseal axis (PATTERSON-KANE et al., 2001). In dogs, the majority of germ cell tumors occur in the ovaries and testicles; few extragenadal neoplasms are reported in the intracranial suprasellar region (HARE et al., 1993; SUMMERS et al., 1995; VALLENTINE et al., 1988) and there is one report of mixed germ cell tumor in the spinal cord (FERREIRA et al., 2003) and

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another one in the eye (PATTERSON-KANE et al., 2001).

A 6-year-old, Airedale Terrier bitch was presented with a history of behavioral changes and abrupt onset of blindness of 7 days duration. Clinical examination revealed apathy, ventromedial unilateral strabismus of the left eye and stiffness of the neck. There was $64\mu\text{g/dl}$ protein in the cerebrospinal fluid. Hematology and blood chemistry profiles were within the normal range. In spite of therapy with corticosteroids and antibiotics, the clinical condition deteriorated and the dog died three days after admission. Gross pathology consisted of $5 \times 4 \times 4\text{cm}$ suprasellar mass which compressed the ventral surface cerebrum from the level of basal ganglia to the mesencephalon (Figure 1A). The cut surface of the mass was soft, red alternated with white areas. The pituitary gland could not be found. No other gross lesions were seen at necropsy.

Tissues were fixed in 10% formalin, embedded in paraffin, sectioned at $5\mu\text{m}$, and stained

with hematoxylin and eosin (HE). Immunohistochemical staining for alpha-fetoprotein (AFP) and cytokeratin were performed in selected slides. Positive controls were fetal liver (for AFP) and skin (for cytokeratin).

Microscopically, compressing and invading the cerebral parenchyma, there was an unencapsulated, expansile neoplasm composed of nests and trabeculae of round to polygonal cells that occasionally surrounded tubules and cysts formed by columnar cells (Figure 1B). All were supported on a fine fibrovascular stroma. The first population had variable distinct borders and moderate amounts of eosinophilic cytoplasm that frequently contained discrete clear vacuoles in a subpopulation somewhat resembling hepatocytes. Rarely, cells contained multiple, eosinophilic, round, $2\text{-}3\mu\text{m}$ diameter secretory granules. Occasionally, columnar cells lining cystic spaces had cilia. Neoplastic cells had oval, vesicular nuclei with 1-2 prominent nucleoli. There was moderate

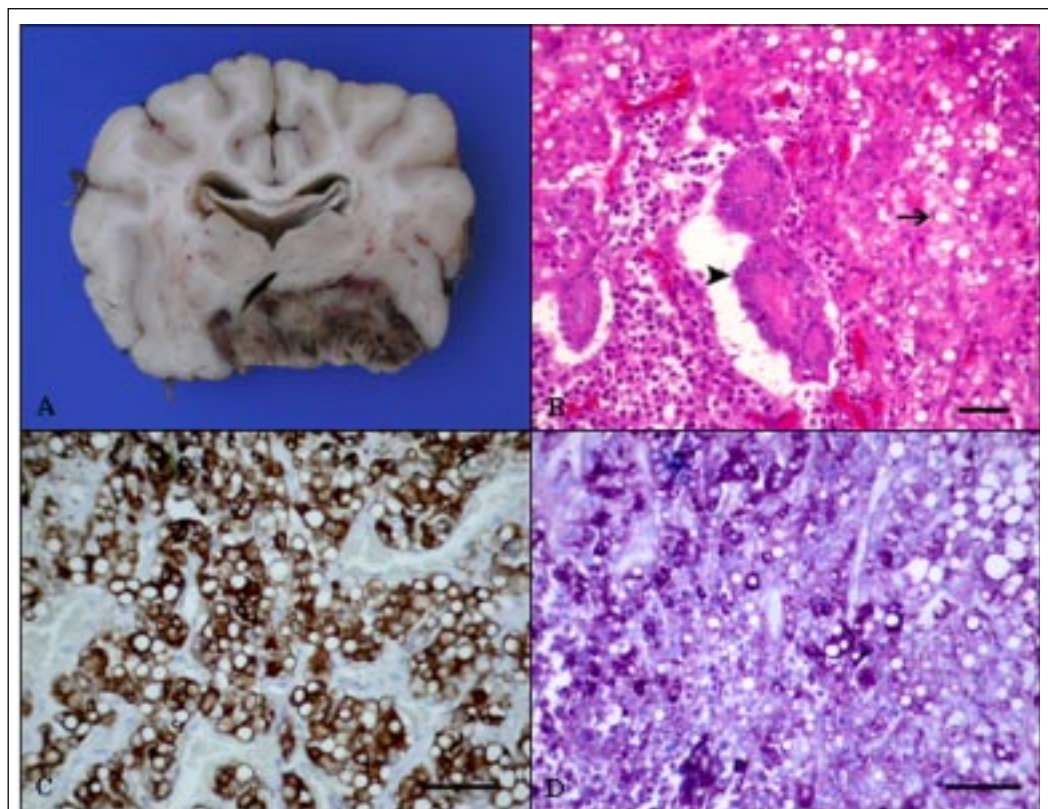


Figure 1 - A. Brain; dog. Suprasellar germ cell tumor. Transverse section at the diencephalon. a variegated brown and white mass has replaced part of the hypothalamus and internal capsule. B. Nests of germ cells and vacuolated hepatoid cells (arrow). There is one acinar structure composed of epithelial cells (arrow head). Hematoxylin-eosin. Bar, $80\mu\text{m}$. C. Positive immunolabeling (brown) of cytokeratin in the cytoplasm of most tumor cells. Hematoxylin counterstain. Bar, $95\mu\text{m}$. D. Immunolabeling (blue) of alpha-fetoprotein in the cytoplasm of some tumor cells. Hematoxylin counterstain. Bar, $95\mu\text{m}$

anisokaryosis and the mitotic rate averages 1-2 per 10 high power fields. Within the neoplasm there were variably sized areas of hemorrhage and necrosis and there were fibrin thrombi. Neoplastic cells were immunopositive for cytokeratin (Kermix) (Figura 1C) and alpha-fetoprotein (Figura 1D).

There is considerable debate on the origin of suprasellar pleomorphic neoplasms. Suprasellar germ cell tumors presumably develop from ectopic germinal epithelium, whereas in humans, craniopharyngiomas presumably derive from remnants of Rathke's pouch (HAWKINS et al., 1985; KOESTNER & HIGGINS, 2002). Recently, it has been proposed that suprasellar pleomorphic neoplasms be classified as suprasellar germ cell tumors rather than craniopharyngiomas based on the following criteria: (1) midline suprasellar location; (2) varying populations of neoplastic cells composed of a mixture of germinatous (seminoma/dysgerminoma-like) areas, hepatoid cells, and areas with intestinal or respiratory epithelial differentiation; and (3) positive staining for alpha-fetoprotein (KOESTNER, 1999; NYSKA et al., 1993). Therefore, in this case, the characteristic histomorphologic features, location, and results of the immunohistochemical stains support the diagnosis of a suprasellar germ cell tumor.

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