Meningioma in cervical spinal cord segment 6 of a dog – a case report

Ciprian Andrei Ober¹, Orit Chai², Joshua Milgram², Cosmin Petru Peștean¹, Cecilia Danciu¹, Teodoru Soare³, Liviu Ioan Oana¹, Marian Taulescu¹

¹University of Agricultural Sciences and Veterinary Medicine, Faculty of Veterinary Medicine, Cluj-Napoca, Romania
²Hebrew University of Jerusalem, Koret School of Veterinary Medicine, Rehovot, Israel
³University of Agricultural Sciences and Veterinary Medicine, Faculty of Veterinary Medicine, Bucharest, Romania

Received November 28, 2017
Accepted August 13, 2018

Abstract

Meningiomas in dogs occur more commonly in the brain than in the cranial spinal cord. Intramedullary spinal cord tumours in dogs are described infrequently and present a diagnostic and therapeutic challenge. A nine-year-old Beagle dog was referred because of tetraparesis of a 20-day duration. The neurological signs were suggestive of a selective lesion involving the cervical spinal cord. Sagittal T2-weighted magnetic resonance imaging of the cervical vertebral column revealed a ventral, well-circumscribed mass within the vertebral canal at the level of cervical segment 6 (C6). A primary neoplasia was considered as probable differential diagnosis. The mass was removed by cervical laminectomy, durotomy and gentle dissections. On the basis of histological and immunohistochemical findings, a diagnosis of transitional meningioma (grade I) was made. Treatment of the meningioma with surgery resulted in a complete recovery, the dog was able to walk 21 days after surgery and had normal walk two months after presentation. Clinicopathologic and treatment data of cranial intraspinal meningiomas have been reported sporadically, but a segment 6 location was not thoroughly described before.

Grading system, canine, tumour

Meningioma is the most common primary nervous system tumour of the spinal cord of the dog (Wright 1985; Drost et al. 1996; Levy et al. 1997; Petersen et al. 2008; Wu et al. 2011). Because of striking similarities between canine and human meningiomas (Kraft et al. 1997), the human World Health Organization (WHO) classification system has been applied to canine meningiomas in the recent years (Sturges et al. 2008; Mandara et al. 2010). Typically, intraspinal meningiomas cause a chronic, progressive myelopathy with mild to moderate spinal pain (Fingeroth et al. 1987; Bell et al. 1992; Vanwinkle et al. 1994; Auger et al. 1996; Forterre et al. 2002; De Bosschere et al. 2003; Dunie-Merigot and Huneault 2006). Middle-aged to older dogs and cats are most commonly affected (Koestner and Higgins 2002; Montoliu et al. 2006). Meningiomas develop from the neoplastic transformation of arachnoid (meningothelial) cells (Kepes 1986; Burger and Scheithauer 2007). Their highly diverse microscopical appearance might be explained by the dual contribution of mesoderm and neural crest to meningeval development (O’Rahilly and Müller 1986), providing the capacity for neoplastic arachnoid cells to undergo mesenchymal and epithelial differentiation (Kepes 1986). While the thoracic spine is the most common location of spinal meningiomas in humans (80%), canine spinal meningiomas are more frequent in the cranial part of the cervical spinal cord (Fingeroth et al. 1987; Levy et al. 1997; Barnhart et al. 2002; Montoliu et al. 2006; Beall et al. 2007; Petersen et al. 2008). Definitive diagnosis is essential and requires biopsy and histopathological examination. A tentative diagnosis is often possible based on tumour characteristics using advanced imaging techniques (José-López et al. 2013).
Typical magnetic resonance imaging characteristics include a contrast-enhancing mass with a broad-based dural attachment and variable signal intensity on precontrast T1- and T2-weighted images (Vanwinkle et al. 1994; Drost et al. 1996; Kippenes et al. 1999; McDonnell et al. 2005). Most histopathologic characteristics of canine meningiomas are strikingly similar to those in humans (Koestner et al. 1999; Kleihues and Cavenee 2000; Sturges et al. 2008). Tumour grading is an important diagnostic criterion in human intracranial meningiomas and has both predictive and prognostic values (Kleihues and Cavenee 2000). Although several reports have described various histologic subtypes in canine spinal cord meningiomas, tumour grading is rarely used (Barnhart et al. 2000; Theon et al. 2000). The aim of this study was to describe the clinical, imaging and pathologic characteristics of a meningioma in a rare location at the level of cervical spinal cord segment 6 in a dog.

Case presentation

A 9-year-old, male intact Beagle was admitted with a 20-day history of right thoracic limb monoparesis that progressed to ambulatory tetraparesis. No traumatic incident was reported by the owners. On neurologic examination there was moderate ambulatory tetraparesis, worse on the right side. Postural reactions were absent in the right thoracic limb and delayed in the other limbs. The rest of the neurological examination was unremarkable. Results of the complete blood count, serum biochemical profile, thoracic radiographs, and abdominal ultrasonography were within normal limits. Survey radiographs of the vertebral column under general anaesthesia did not reveal abnormalities. Sagittal T2-weighted magnetic resonance imaging of the cervical vertebral column revealed a ventral, well-circumscribed mass in the vertebral canal over the 6th cervical vertebra (C6) (Plate III, Fig. 1A,B). The spinal cord was severely compressed at this level by the mass. Cerebrospinal fluid analysis results were unremarkable.

After premedication with diazepam (Terapia SA, Romania) (0.2 mg/kg intravenously), anaesthesia was induced with propofol (Lipuro 1%, Braun Melsungen AG, Germany) (4 mg/kg intravenously) and maintained with isoflurane (Forane, Abbots Laboratories, UK) in oxygen. The dog was positioned in sternal recumbency with the head gently flexed in a neutral position. A dorsal approach to the caudal cervical spine was performed. The multifidus musculature on the spinous processes was elevated with a periosteal elevator to expose the dorsal bony lamina. The spinous processes and the yellow ligament were removed with a rongeur to the level of the dorsal bony lamina, which was removed with a high speed drill. The dura mater was incised using a No. 11 blade and fine tenotomy scissors. The mass and compressed spinal cord were observed (Plate III, Fig. 1C). A bipolar cautery, loupes and copious irrigation with suction to preserve visibility were used. Tumour removal was done by gentle blunt dissection, creating a plane between normal tissue and the mass (Fig. 1C, D). The dural incision was not sutured. The muscles, subcutaneous tissues, and skin incision were apposed in layers. Pain management was achieved by treatment with tramadol (Ozone Laboratories Ltd, UK) (4 mg/kg orally), and carprofen (Rimadyl; Zoetis, USA) (2 mg/kg, orally). A first generation cephalosporin, cefadroxil (Antibiotice, Romania) (22 mg/kg every 8 h, orally) was also used for 10 days postoperatively.

The dog’s owner provided a written, informed consent for the clinical assessment, diagnostic work-up, treatment and follow-up of their pet and for the inclusion of their pet’s information in this manuscript for publication.

Histopathological and immunohistochemical examination

For histopathological examination, tissue samples from the neoplasm were fixed in 10% phosphate buffered formalin for 24 h, trimmed and embedded in paraffin. Sections
were cut at 5 μm and stained with haematoxylin and eosin (H&E). Immunohistochemistry was performed to detect the origin of the neoplastic cell population. A panel of three immunohistochemical markers that included vimentin, pancytokeratin and S100 protein was selected (Table 1).

Table 1. Antibodies used for immunohistochemistry.

<table>
<thead>
<tr>
<th>Antibody</th>
<th>Source</th>
<th>Antigen retrieval</th>
<th>Detection method</th>
</tr>
</thead>
<tbody>
<tr>
<td>Vimentin</td>
<td>Dako</td>
<td>Dako</td>
<td>EnVision</td>
</tr>
<tr>
<td>V9 Clone</td>
<td>Dako</td>
<td>FLEX TRS</td>
<td>FLEX/HRP</td>
</tr>
<tr>
<td>M0725</td>
<td>Low ph</td>
<td>SM 802</td>
<td></td>
</tr>
<tr>
<td>S-100</td>
<td>Dako</td>
<td>Dako</td>
<td>EnVision</td>
</tr>
<tr>
<td>Z0311 Polyclonal rabbit</td>
<td>FLEX TRS</td>
<td>FLEX/HRP</td>
<td></td>
</tr>
<tr>
<td></td>
<td>High ph</td>
<td>SM 802</td>
<td></td>
</tr>
<tr>
<td>Pan-cytokeratin</td>
<td>Dako</td>
<td>Dako</td>
<td>EnVision</td>
</tr>
<tr>
<td>AE1/AE3</td>
<td>Dako</td>
<td>FLEX TRS</td>
<td>FLEX/HRP</td>
</tr>
<tr>
<td>M3515 Monoclonal mouse</td>
<td>Low ph</td>
<td>SM 802</td>
<td></td>
</tr>
</tbody>
</table>

Histologically, the neoplasm was well delimited, partially encapsulated and expansile, composed of two types of cell population. The first population consisted of uniform meningothelial cells arranged in lobules surrounded by thin collagenous stroma (Plate III, Fig. 2A). The second population was composed of numerous spindle-shaped cells, resembling fibroblast and forming intersecting streams embedded in a moderate amount of pale eosinophilic collagen-rich matrix. Multifocally, the neoplastic cells formed whorls or variable sized round structures of concentric layers. The latter structures were often hyalinized and slightly calcified, interpreted as psammoma bodies (Fig. 2B). The presumptive diagnosis of mixed (transitional) meningioma was made. Expression of S-100 (Fig. 2C) and vimentin (Fig. 2D) protein antibodies was represented by an intense labelling of the cell nuclei of the neoplastic cells. Pan-cytokeratin immunoexpression was diffusely negative. These findings demonstrate the mixed origin of the neoplastic cells and were compatible with the diagnosis of a transitional meningioma.

**Discussion**

Reports of canine meningiomas in the caudal part of the cervical spinal canal are scarce. The most meningiomas affect the spinal cord cranial to vertebra C3 (Fingeroth et al. 1987; Vanwinkle et al. 1994; McDonnell et al. 2005; Petersen et al. 2008). Of the 53 reported cases of spinal cord meningiomas for which location was mentioned (Zaki et al. 1975; Gilmore 1983; Raskin 1984; Fingeroth et al. 1987; Vanwinkle et al. 1994; Siegel et al. 1996; Levy et al. 1997; Kippenes et al. 1999; Yeomans 2000; Forterre et al. 2002; Barnhart et al. 2002; De Bosschere et al. 2003; McDonell et al. 2005; Dunie-Merigot and Huneault 2006; Petersen et al. 2008), 67% were cervical, 4% thoracic, 23% lumbar, and 6% were multilocular. Tumour position was lateral in 32%, posterolateral in 10%, posterior in 3% and anterior in 13%. A meningothelial cystic mass was found in a dog at the C5-C6 level (José-López et al. 2013).

A diagnosis of transitional meningioma can be confirmed by histopathological and immunohistochemical examination of tissue obtained by surgical excision or at necropsy.
Recently, for dogs, these tumours have been graded on the basis of the mitotic count and histological criteria for malignancy, according to the WHO system for human beings (Kleihues et al. 2002), as grade I (benign), grade II (atypical), and grade III (anaplastic) (Petersen et al. 2008; Sturges et al. 2008). Most meningiomas affecting the canine spinal cord are grades I or II, while grade III tumours are rare (Petersen et al. 2008). Grade I tumours appear to be more likely to develop in older dogs and are more likely to be located in the cervical region (Petersen et al. 2008). There is no apparent correlation between the grade of tumour and imaging characteristics or the long-term outcome (Petersen et al. 2008). Cerebrospinal fluid analysis may be normal or may reveal nonspecific abnormalities and neoplastic cells are rarely identified. Cerebrospinal fluid analysis was performed in our case but no neoplastic cells were identified.

Aggressive behaviour of some spinal meningiomas has been observed. Thus, in a study of 13 spinal meningiomas (Petersen et al. 2008) four invasive tumours were identified. Diagnoses of osseous-associated cervical spondylomyelopathy or meningioma were considered the strongest variables to predict early post-operative neurological deterioration, associated with higher (more severely affected) neurological grade before surgery and longer surgery time.

Reported treatments for canine intraspinal meningiomas consist of medical management and cytoreductive surgery with or without radiation therapy (Raskin 1984; Fingeroth et al. 1987; Bell et al. 1992; Vanwinkle et al. 1994; Auger et al. 1996; Siegel et al. 1996; Levy et al. 1997; Forterre et al. 2002; De Bosschere et al. 2003; Dunie-Merigot and Huneault 2006; McDonnell et al. 2005). Survival times after surgery vary from 4 to 47 months (Fingeroth et al. 1987; Bell et al. 1992; Vanwinkle et al. 1994; Auger et al. 1996; Levy et al. 1997; Forterre et al. 2002; De Bosschere et al. 2003; Dunie-Merigot and Huneault 2006). The effect of adjunctive radiation therapy on the survival time is unclear, because there are only a few reports describing its use (Bell et al. 1992; Siegel et al. 1996; McDonnell et al. 2005).

Surgical gross total tumour removal of vertebral tumours improves the quality of life (Besalti et al. 2016). No surgical complications were observed in our case. The duration of hospitalisation was seven days. Paresis and absent pelvic limb postural reactions were still present in the first five days but the dog was able to rise and make a few steps with assistance after seven days post operation. Noticeable ataxia with some limb deficits were observed by the owner in the first month after discharge, but no cervical hyperaesthesia was noticed. Progressive neurological improvement was reported by the owner with a complete resolution of neurological signs observed two months after surgery at a follow-up at our clinic. We did not perform any restaging of this tumour.

Acknowledgements

The authors gratefully acknowledge Cornel CĂTOI, Rector of University of Agricultural Sciences and Veterinary Medicine, Cluj-Napoca, and the dog’s owner for their support of this work.

References

Fig. 1 Sagittal (A) and axial (B) T2-weighted magnetic resonance imaging of the cervical vertebral column. A single intra-axial, well-circumscribed lesion involving the right ventrolateral aspect of the spinal cord over the C6 is observed. C) Intraoperative aspect (note the cord compression, blue arrow). D) Gross appearance of the mass after excision.

Fig. 2. Histological and immunohistochemical features of the transitional meningioma (grade I). A) The photomicrophotograph showing a lobular arrangement of the neoplastic meningothelial cells admixed with numerous spindle-like cells. Haematoxylin and eosin stain. B) The neoplastic cells form multifocal round structures of concentric layers, which are often hyalinized and slightly calcified (psammoma bodies). C and D) Positive immunohistochemical expression of S-100 protein (C) and vimentin (D) antibodies characterized by an intense labelling of the cytoplasm of the neoplastic cells. Streptavidin–biotin complex, DAB–horseradish peroxidase, Mayer’s haematoxylin counterstain.