

Case report

# NASAL TRANSMISSIBLE VENEREAL TUMOR IN A JACK RUSSEL TERRIER BITCH

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A two-year old Jack Russell Terrier female with left-sided epistaxis was brought to the Clinic of the Bucharest Faculty of Veterinary Medicine. Endoscopy of the left nasal cavity revealed a cauliflower-like mass. Cytological and histopathological features were specific to canine transmissible venereal tumour (CTVT). Tumor tissue showed positive immunoreactivity to anti-vimentin monoclonal antibody (90%) and Ki67 (35%) and samples were negative for anti-cytokeratin monoclonal antibody. Chemotherapy with vincristine sulphate at a dose of 0.025 mg/kg i.v./week for 6 administrations was successful. This paper presents a primary form of nasal CTVT in a Jack Russell Terrier female.

Key words: canine transmissible venereal tumor, complex diagnosis, nasal localisation

#### INTRODUCTION

Canine transmissible venereal tumour (CTVT) is a malignant tumor transmitted naturally through a transplant of tumoral cells [1,2]. The main localisation is genital, CTVT affecting the vulva / vaginal vestibule of females and the penis in males [2]. The tumor can also present an extragenital localisation, appearing in the mouth, nasal cavities, eye and skin [1,2]. Tumor development through an implantation of tumoral cells in the mouth or nasal cavities is favoured by the particular behaviour of animals during the mating period (sniffing, licking and biting). The pattern of this neoplasm is a round cell sarcoma. Hematoxylin and eosin staining is observed in round cells arranged or grouped in cords interspersed with a rich connective stroma [2,3]. In this paper, we present a case of CTVT with intranasal localisation in a Jack Russell Terrier bitch assessed by several methods of examination and monitored for over 6 months.

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#### **CASE PRESENTATION**

A two-year old Jack Russell Terrier female was brought to the Clinic of the Bucharest Faculty of Veterinary Medicine. The female had been suffering from left sided epistaxis for five months. Radiological examination was performed with a Röntgen device (Philips Optimus Bucky Diagnost). Endoscopy of the nasal cavities was performed using a Riester GmbH/3920 endoscope. For anaesthesia, the bitch was premedicated with 10 mg/kg medetomidine (Domitor®, Pfizer) and 0.2 mg/kg im butorphanol (Butomidor<sup>®</sup>, Richter-Pharma). Anaesthesia was mantained with 1.5% isoflurane (Anesteran<sup>®</sup>, Rompharm Co.) on a Dragger Fabius anaesthetic machine. Cytology was performed on nasal secretions (from both nasal cavities) and fragments of tumour tissue harvested during endoscopy. Slides were stained with May Grunwlad Giemsa stain. For the histopathological examination, samples were fixed in a 10% formalin solution, embedded in paraffin, cut at 5 µm and stained with haematoxylin and eosin. Immunohistochemistry (IHC) was performed on 5 µm thick sections from 10% formalin-fixed paraffin-embedded tissues using the Streptavidin-Biotin (SABC) /Horseradish Peroxidase (HRP) method (StreptABC, Dako, Denmark). Monoclonal antibodies were used for this examination: anti-vimentin, anti-cytokeratin and anti-Ki67 antigen. Therapy consisted of six intravenous administrations of vincristine sulphate (Sindovin, Sindan Pharma Romania) at a dose of 0.025 mg/kg/week. Epistaxis is common in nasal forms of CTVT (Figure 1) [4-6]. Early haemorrhage is explained by particularities in the consistency and structure of the mass; CTVT is a well vascularised friable tumor par excellence [5,7]. The location presented in this paper has been previously reported by other authors [5-7]. The nasal form of CTVT has been diagnosed both in females and males. Its incidence is related to coital behaviour (licking, biting, sniffing). Radiography of the skull by rostro-caudal incidence revealed an area of increased radiopacity in the left frontal sinus. Increased radiopacity of the frontal sinus occurs in: sinusitis (with a bacterial, fungal or allergic aetiology), blockage of the sinus drainage with mucus accumulation and malignant tumours [8]. We believe the radiological imaging is not specific for the final diagnosis of CTVT, but this investigation is extremely important to assess the extent of lesions and bone invasion (a factor which leads to a poor prognosis). During endoscopy in the left nasal cavity the following lesions were found: abundant capillary bleeding, mucosal erosions, ulcers and multiple cauliflower-like friable lesions (Figure 2). Bleeding was more discreet in the right nasal cavity as compared to the left one. The rhinoscopy results are common for the following pathological processes: rhinitis and neoplasia. For the nasal localisation of CTVT, the gross lesions observed during rhinoscopy were described by Papazoglou et al. [4,5].

Groups of round cells with vacuolated basophilic cytoplasm, chromatin with granular appearance and 1-2 eccentrically located nucleoli were observed at the cytopathological examination. According to cytological criteria described by Anne Santos Amaral [9], we can state that the observed population was dominated by lymphocytoid type cells (Figure 3). The histopathological examination revealed round cells arranged or grouped in cords supported by a rich connective stroma. The neoplastic cells arrangement shows an arborescent layout (specific to CTVT), with a rich capillary network (Figure 4). According to the histopathological appearance, CTVT falls into the pattern of a round cell sarcoma [3,10].

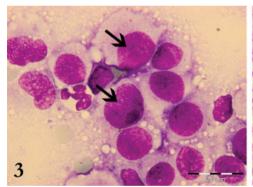




Figure 1. Clinical aspect, left-sided epistaxis



Figure 2. Endoscopy of the left nasal cavity. Common nasal meatus mucosa presents ulcerative lesions as well as a tumor mass (arrow)



granular chromatin (arrow tail) and eccentric stroma (arrow tail) nucleolus and cytoplasm with vacuoles

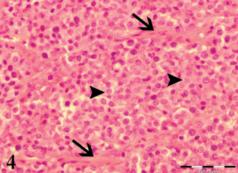


Figure 3. CTVT cytological appearance (May Figure 4. CTVT histopathologic aspect (HE), Grunwald Giemsa), 1000x, bar 20 µm. Round 400x, bar 50 µm. Round cells arranged in cells with anisocytosis and anisokariosis with rows (arrowhead) supported by a conjunctive

Samples were negative for immunostaining with AMC anti-cytokeratin (clone MNF116, Dako Glostrup Denmark). The tumor showed an intense immunoreactivity (90%) to AMC anti-vimentin (clone V9, Dako Glostrup Denmark). The immunohistochemical research carried out by Mozos (1996) and Marchal (1997) showed the reactivity of vimentin to be over 90% for CTVT and canine cutaneous histiocytoma [11,12].

In our study there was an intense immunostaining of tumoral cells with anti-vimentin AMC clone V9 (Figure 5), which is consistent with literature reports. In the current study, immunostaining with Ki-67 antigen, clone MIB (Dako Glostrup, Denmark), showed a tumor proliferation index of 35% (Figure 6).

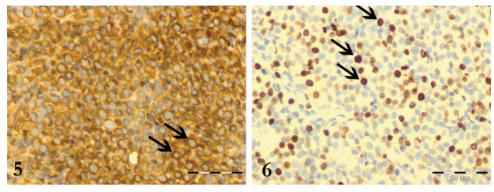


Figure 5. Vimentine positive (90% of the tumor cell cytoplasm – arrow) in CTVT nasal location (imunohistochemistry 400x, bar 50  $\mu$ m)

Figure 6. Ki-67 positive (35% of tumor cell nuclei – arrow) in CTVT nasal location (imunohistochemistry, 400x, bar 50  $\mu$ m)

Ki-67 antigen is a marker of cell cycle and cellular proliferation used to estimate the coefficient of proliferation in a cell population. Research has shown that the Ki-67 antigen plays an important role in assessing malignancy [13].

Therapy consisted of 6 doses of 0.025 mg/kg vincristine sulphate administered every 7 days. After the first two doses, relevant clinical signs (epistaxis and sneezing) began to subside. At the last check-up (30 days after the start of therapy), the animal did not present any clinical signs.

Six months after the treatment was completed the owner confirmed that the patient was in a good condition (it showed no clinical signs of disease) and that the diagnosis and the therapeutic approach had been successful. When comparing our results to existing data, we see similar outcomes [14].

The novelty of this work is the report of a nasal primary form of CTVT in a bitch. In the previous cases this localisation was most frequently diagnosed in males. There is no gender proneness for CTVT, the tumor occurring in both females and males. Nasal localisation of CTVT is more frequent in males due to behavioural features: licking, sniffing and biting of the genital region as well as of other body parts [15,16].

In the area where the animal presented in our study lives (the Bucharest metropolitan area) CTVT is frequently diagnosed in stray dogs, which provide the tumour with a continuous natural reservoir. We believe that in this case a stray dog may have been the source of contamination with tumoral cells.

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## NAZALNI TRANSMISIVNI VENERIČNI TUMOR KOD KUJE RASE JACK RUSSEL

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Na klinici Fakulteta veterinarske medicine u Bukureštu, prezentovana je dve godine stara kuja rase Jack Russel sa epistaks krvavljenjem iz leve nozdrve. Endoskopijom leve nosne šupljine ustanovljena je masa karfiolaste strukture. Citološke i histopatološke karakteristike su odgovarale karakteristikama specifičnim za transmisivni venerični tumor (TVT) pasa. Tumor je bio imunohistohemijski pozitivan na anti-vimentin monoklonsko antelo (90%) kao i na Ki67 (35%), a negativan na anti-citokeratin monoklonsko antitelo. Hemoterapija sa vinikristin sulfatom, u dozi od 0.025 mg/kg telesne mase datog intravenski jedanput nedeljno, u trajanju od ukupno 6 nedelja, bila je uspešna. U radu se prikazuje primarna forma nazalnog TVT kod ženke rase Jack Russel terijera.