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Occurrence and pathology of cutaneous tumours in dogs[#]

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Abstract

The current research work was undertaken to study the occurrence and pathology of cutaneous tumours in dogs and their age, breed, location and gender-wise distribution. A total of 33 samples suspected of cutaneous tumours were collected from the cases presented to Teaching Veterinary Clinical Complex (TVCC), Pookode, Wayanad, Kerala during a period of 12 months from June 2022 to June 2023. Based on histopathology, most of the tumours were diagnosed as benign. Majority of the tumours were of mesenchymal origin. Among different histological types, highest occurrence was that of squamous cell carcinoma (SCC) and trichoblastoma followed by histiocytoma, lipoma, mast cell tumour, hepatoid gland adenocarcinoma and haemangioma. Although, the incidence was higher among males and among the dogs belonging to the age group of 5-10 years, it was not statistically significant. Among different breeds, highest occurrence was observed in Labrador Retriever followed by German Shepherd. The study also warrants large scale epidemiological studies to identify the risk factors and to unravel the etiopathogenesis of canine cutaneous neoplasms.

Keywords: Cutaneous tumours, dogs, occurrence, histopathology

Cutaneous neoplasms in dogs continue to be an important disease in pet animal practice and the most leading cause of death in dogs (Rafalko *et al.*, 2023). Dogs are twice as likely as humans to develop tumours (Rungsipipat *et al.*, 2003). As being companion animals, sharing a common environment with humans, they can acquire a variety of spontaneous tumours during their

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lifetime similar to humans (Gamlem et al., 2008). Tumours of skin are the most and second-most common tumours reported in male and female dogs, respectively accounting for approximately one-third of all tumours encountered in dogs (Graf et al., 2018). These tumors can be benign or malignant and may arise from various cell types present in the skin and can affect dogs of all ages, breeds and sizes. The high incidence of development of tumours in skin could be due to continuous exposure of skin to various physical, chemical and other environmental factors (Subapriya et al., 2021). This may also be due to early detection of these cutaneous lesions. The incidence of cutaneous tumors in canines has been on the rise in recent years, making it an important area of research and veterinary attention. Several surveys undertaken in different countries have revealed geographical variations in the occurrence and type of skin tumours that develop in dogs. The discrepancies are attributable to geographical differences in environmental carcinogen exposure and breed popularity (Mukaratirwa et al., 2005). This article aims to provide an overview of the occurrence and clinicopathological features of cutaneous neoplasms in dogs during the period of June 2022 to June 2023 in and around Wayanad.

Materials and methods

The present study involved collection of suspected cutaneous neoplastic samples from the canine cases presented to Teaching Veterinary Clinical Complex, Pookode. A total of 33 samples were collected during a period of 12 months from June 2022 to June 2023. Clinical data of cases such as age, species, breed, sex, location, shape and size were recorded. The samples were fixed in 10 per cent neutral buffered formalin (10% NBF), processed and stained using routine haematoxylin and eosin (H&E) staining method for histopathological examination. Chi-square test with Yate's correction and Fischer's exact tests were used to find out the relationship between tumour occurrence and other categorical variables using GraphPad Prism v.9.

Results and discussion

Tumours in the present study were classified into epithelial/ melanocytic origin

and mesenchymal origin based on World Health Organisation (WHO) classification. Among all the tumours of cutaneous origin, five cases (15.15 per cent) each of squamous cell carcinoma (SCC) and trichoblastoma were observed. It was followed by four cases (12.12 per cent) of histiocytoma, three cases (9.09 per cent) of lipoma, mast cell tumour, hepatoid gland adenocarcinoma, haemangioma and two cases (6.06 per cent) of melanoma, haemangiopericytoma, angiofibroma and one case (3.03 per cent) of liposarcoma. Among all tumours, the occurrence of squamous cell carcinoma and trichoblastoma were found to be more in the present study. This result was found in agreement with observations made by Lafta and Alaboddy (2020) in their study on incidence of cutaneous tumours in dogs.

Trichoblastoma

Five cases of trichoblastoma were examined and among these, two cases revealed a solid pattern with mild to moderately dense fibrous stroma dividing neoplastic cells into multiple islands (Fig. 1). One case showed a medusoid pattern in which multiple long, thin cords of cells appeared to be arising from a central core (Fig. 2) and one case had a granular cell type pattern. Ribbon type of trichoblastoma was identified in one case in which the neoplastic cells were arranged in two or three cell thick cords. The neoplastic cells had scanty basophilic cytoplasm with indistinct cell borders, round to oval nucleus and hyperchromatic inconspicuous nucleoli with a palisading arrangement. These findings were in accordance with the reports of Goldschmidt and Goldschmidt (2017).

Squamous cell carcinoma

Out of the five cases of squamous cell carcinoma (SCC) examined in the study, three cases were well differentiated and two cases were moderately differentiated. The welldifferentiated neoplasm was characterised by neoplastic squamous epithelial cells invading the dermis and subcutaneous tissue along with distinct keratin pearl formation, the cells were arranged in concentric layers with central core of keratin (Fig. 3). The neoplastic cells were large and ovoid to polyhedral with pale eosinophilic abundant cytoplasm and the nuclei were round to oval, vesicular and centrally located with single prominent nucleoli and frequent mitotic figures. These observations were in agreement with the findings of Kashyap *et al.* (2013). The moderately differentiated neoplasms were characterised by multiple variably sized islands of neoplastic epithelial cells with fewer keratin pearls (Fig. 4).

Hepatoid gland adenocarcinoma

Histopathological examination of hepatoid gland adenocarcinoma revealed multiple lobules of hepatocytes like neoplastic cells that were separated by variable amounts of fibrovascular stroma. The reserve cells were characterised by round, small cells with basophilic cytoplasm and hyperchromatic nuclei that were arranged at the periphery of the lobules. The neoplastic cells were pleomorphic with abundant, highly eosinophilic cytoplasm, which was sometimes vacuolated and granular. The nuclei were pleomorphic, large and located centrally with prominent nucleoli and occasional mitotic figures were evident (Fig. 5). These findings were in concordance with observations of Yumusak et al. (2016).

Melanoma

Histopathological examination of two cases of digital melanoma revealed melanocytic cells which were obscured by melanin pigment and were arranged in nests and sheets separated by thin fibrous stroma in dermis and dermo-epidermal junction (Fig. 6). The melanocytic cells were large round to polygonal with distinct cell borders containing oval vesicular nuclei with multiple prominent nucleoli. These findings were in accordance with reports of Goldschmidt and Goldschmidt (2017).

Lipoma

All three cases of lipoma examined in the present study were histologically characterised by neoplastic adipocytes with vacuolated cytoplasm that pushed the nuclei into periphery and appeared flattened along the cell membrane (Fig. 7). A thin to thick fibrous tissue stroma separated adipocytes into multiple lobules. These findings were similar to the reports by Mathew *et al*. (2020).

Angiofibroma

Angiofibroma, microscopically, appeared as a moderately cellular mass composed of irregular group of blood vessels within a connective tissue matrix. Higher magnification revealed spindle or stellateshaped fibroblasts with oval to elongated nucleus (Fig. 8). Present findings were in accordance with the observations made by Kluthcovsky *et al.* (2021).

Haemangioma

Histopathological examination of haemangioma revealed well-circumscribed cavernous type characterised by variably sized, widely dilated vascular channels filled with erythrocytes separated by varying amount of fibrous connective tissue stroma (Fig. 9). These channels were lined by a single layer of endothelial cells containing inconspicuous nuclei. These findings were in agreement with Lather *et al.* (2014).

Haemangiopericytoma

Histopathological examination of haemangiopericytoma revealed highly cellular mass composed of spindle cells arranged concentrically around capillaries, like whorls in a collagenous stroma (Fig. 10). Neoplastic spindle cells were having moderate eosinophilic cytoplasm with vesicular nuclei and inconspicuous nucleoli. Our findings were in agreement with reports of Simeonov *et al.* (2011).

Mast cell tumour

Three cases of mast cell tumours (MCT) were documented in the study. Based on the cellular and nuclear morphology, architecture and cellularity, two cases were graded as grade 1 (low grade) and one was grade 2 (intermediate grade) according to Patnaik three-tier grading system. Low grade MCT was characterised by cells arranged in rows separated by mature collagen bundles (Fig. 11). The cells were round with distinct moderate cytoplasm and mostly eccentrically placed nucleus, which was round with condensed chromatin. Intermediate grade MCT was characterised by highly cellular mass with mast cells arranged in rows and sheets with bright eosinophilic bands of collagen along the course of neoplastic mast cells (Fig. 12). The moderate pale eosinophilic cytoplasm contained fine granules and round to oval, centrally or eccentrically placed nucleus.

Histiocytoma

Histopathologically, in all four cases of histiocytoma, the cells were arranged in sheets separated by thin band of collagen in the dermis replacing most of the adnexal structures. The cells were round with varying amount of pale eosinophilic, sometimes vacuolated cytoplasm and round to oval to intended nucleus was observed (Fig. 13). These findings were in accordance with observations of Goldschmidt and Goldschmidt (2017).

Liposarcoma

Histopathologically liposarcoma was characterised by round to large polyhedral cells arranged in sheets and lobules by intervening connective tissue septa. Smaller portion of cells had single large fat vacuole with peripheral nucleus. Some cells were foamy with variably sized vacuoles in the cytoplasm (Fig. 14). These findings were in agreement with Avallone *et al.* (2016).

Incidence of tumours

Out of 33 cutaneous tumour cases, 22 tumours (66.67 per cent) were diagnosed as benign and remaining 11 tumours (33.33 per cent) were diagnosed as malignant, based on histopathological examination. The increased incidence of benign tumours was in accordance with Graf *et al.* (2018) and Martins *et al.* (2022) who reported 57.52 per cent and 62.90 per cent of tumours as benign in their study. However, this observation was not in agreement with Sharma *et al.* (2018) and Karnik *et al.* (2020) who reported increased incidence of malignant tumours in their studies. On gross examination, majority of tumours had round to oval shape with size varied from 1 cm

to 14 cm. Majority of the tumours were soft to firm in consistency and greyish-white colour on cut section. Most cases of squamous cell carcinoma, mast cell tumour, trichoblastoma and hepatoid gland adenocarcinoma were ulcerated. Histopathological examination revealed a higher occurrence of mesenchymal tumours (54.54 per cent). This observation was in consistent with reports of Machado *et al.* (2018) and Kok *et al.* (2019).

In the present study, the highest risk of occurrence of cutaneous tumours was seen in the age group of 5-10 years (42.42 per cent) followed by in less than five years (39.39 per cent) and those in the age group of greater than 10 years (18.18 per cent). Even though more number of tumours were observed in age group of 5-10 years, there was no significant difference in the occurrence of cutaneous tumours (p>0.05) between different age groups. Senthil et al. (2020) observed higher occurrence in the age group of 5-10 years (53.80 per cent) followed by >10 years (23.30 per cent) and below five years (23.10 per cent); whereas, Bhanderi et al. (2022) observed highest frequency in <5 years (39.34 per cent) followed by 5-10 years (37.71 per cent) and >10 years (22.95 per cent). The variation in age susceptibility to the disease may be due to multiple factors including environmental factors like exposure to carcinogens and also due to breed variation in longevity and susceptibility to disease in the various reports.

The occurrence of cutaneous neoplasm was observed more in males (57.58 per cent) than in females (42.42 per cent). This result was in accordance with the observations recorded by Nair et al. (2021) and Subapriya et al. (2021) who reported higher incidence of skin neoplasm in male dogs than females. Though the occurrence of tumours was more in males, there was no significant difference in tumour occurrence (p>0.05) between males and females, which is in agreement with the study of Mathew et al. (2020). Among the presented cases, hepatoid gland adenocarcinoma was found only in male dogs and all other tumours were observed both in male and female dogs. This was in accordance with the reports of Shabeeba et al. (2021) and Subapriya et al. (2021). This might be due to the normal



x100)

multiple islands of basal epithelial cells surrounded by moderate connective tissue stroma (H&E, x100)





Fig. 3. Well-differentiated SCC-Multiple islands separated by fibrous tissue stroma with keratin pearl formation (H&E, x40)



Fig. 4. Moderately differentiated SCC- Cord like arrangement of epithelial cells with keratinisation (H&E, x100)



Fig. 5. Hepatoid gland adenocarcinoma-Cells with granular eosinophilic cytoplasm, anisokaryosis anisocytosis, (H&E, x400)



Fig. 6. Melanoma- Polyhedral cells with eosinophilic cytoplasm, vesicular nucleus with multiple prominent nucleoli (H&E, x400)



7. Lipoma- Closely packed Fig. variably sized round to polyhedral cells with single vacuolated cytoplasm (H&E, x100)

Fig. 8. Angiofibroma- Multiple narrow vessels surrounded by spindle shaped fibroblasts (H&E, x400)

Fig. 9. Haemangioma- Variably sized vascular channels separated by connective tissue stroma (H&E, x40)

expression of androgen receptors in the canine hepatoid gland cells and the association of hepatoid tumours with the male sex hormone androgen as stated by Kaldrymidou et al. (2002).

The occurrence of cutaneous neoplastic conditions was observed more in Labrador Retriever (33.33 per cent; 11/33) followed by German shepherd (18.18 per cent; 6/33), Rottweiler and Boxer (12.12 per cent each; 4/33), spitz and non-descript dogs (9.09 per cent each; 3/33). Similarly, highest incidence of tumours in Labrador dogs was recorded

by Karnik et al. (2020) and Nair et al. (2021). A higher incidence of tumours was noticed in purebred dogs than non-descript dogs. This was in concordance with the observations recorded by Hemanth et al. (2015) and Vascellari et al. (2016). The breed predilection noted in the present study, though cannot be generalised, could partly be due to difference in risk associated with various predisposing alleles in the different dog breeds and can also be due to specific breed popularity in an area.

Out of 33 tumour samples examined in the current study, highest occurrence of

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Fia. Spindle cells arranged in whorls mast cells into the dermis arranged around capillaries (H&E, x400)

10. Haemangiopericytoma- Fig. 11. MCT- Grade I- Infiltration of in rows and sheets separating collagen bundles (H&E, x100)

Fig. 12. MCT- Grade II-Round to polyhedral neoplastic cells with moderate pale pink cytoplasm containing fine granules (H&E, x400)



Fig. 13. Histiocytoma- Cells arranged like sheet with moderate eosinophilic cytoplasm and round to oval to intended hyperchromatic nuclei (H&E, x400)

tumours was reported in trunk (39.39 per cent), followed by perianal and head (15.15 per cent each) and 0.09 per cent each on limb and digit. Two cases each was reported from tail and neck. Our findings on location wise occurrence of skin neoplasms differed with the observations of Subapriya et al. (2021) who recorded 33.13 per cent of neoplasms on limbs followed by 24.30 per cent on trunk in a study of a cutaneous tumours. However, Chikweto et al. (2011) reported a high incidence of skin tumours on trunk as reported in the present study. This could be due to exposure of this vast area to the environmental carcinogens including prolonged solar exposure.

Conclusion

The study was conducted to record the occurrence and clinicopathological features of canine skin neoplasms. The current study identified 33 cases of cutaneous tumours in dogs and histopathological examination confirmed most of the neoplasms as benign and



Fig. 14. Liposarcoma- Cells with indistinct cell borders, several small vacuoles, adipocytes with multiple round nuclei, anisocytosis, anisokaryosis (H&E, x400)

also helped in the classification of the tumours different histotypes. Epidemiological into predilection factors such as age, breed, gender and location were recorded. The data recorded in the present study would be a beneficial reference for the veterinary clinician to make a preliminary diagnosis of cutaneous tumours. The study also warrants large scale epidemiological studies to identify the risk factors and understand etiopathogenesis of canine cutaneous neoplasms, which will give new insights in to the diagnosis, treatment and even prevention of this disease.

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Conflicts of interest

There were no conflicts of interest reported by the authors.

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