+ + + + + + CREATING CLARITY

One lump or two? Clinical approach to mammary tumours in canines



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Disclosure

- I am an IDEXX employee
- Legal disclaimer: The information contained herein is intended to provide general guidance only. As with any diagnosis or treatment, you should use clinical discretion with each patient based on a complete evaluation of the patient, including history, physical presentation, and complete laboratory data. With respect to any drug therapy or monitoring program, you should refer to product inserts for a complete description of dosages, indications, interactions, and cautions. Diagnosis and treatment decisions are the ultimate responsibility of the primary care veterinarian.



Purpose

- Answer common questions received from vets about histology reports with a diagnosis of canine **malignant** mammary tumor
- Provide practical information to plan approach to dogs with mammary tumors prior to surgical removal



Canine mammary tumours

Outline

- Clinical features
- Prognostic factors
- Diagnosis
 - Cytology
 - Histology & risk of recurrence/metastasis
 - Histological grading
 - Specific tumor types
- Surgical approach / recurrence
- Metastasis: LN removal / chemotherapy
- Prognostic role of spaying after diagnosis



Clinical features

Can I guess if a mammary tumor is malignant or benign prior to surgery?

- History: intact/neutered, timing of lesions
- Clinical features
 - Size (>3cm), growth speed and invasiveness
 - Draining lymph node status
 - Appearance of surrounding skin
 - Multiple lesions?
- Staging if suspected malignant
 - Lungs / draining lymph node(s) assessment



⁽Pictures courtesy of Juan Borrego, DACVIM Oncology)

(Gunnes et al. 2017; Gedon et al. 2021)

CLINICAL PRONOSTIC FACTORS

Age - high prevalence of malignant neoplasia

Tissular invasiveness, diffuse growth, skin involvement

Ulceration, rapid growth

Size >3 cm. (T)

Lymph node metastasis (N)

Distant metastasis (M)



TNM Staging system

(Sorenmo et al. 2020)

(Picture courtesy of Juan Borrego, DACVIM Oncology)



Diagnosis: Is cytology of a mammary mass useful?

- Rule in/out:
 - Inflammatory aetiology
 - Malignant mammary tumours (only rule in)
 - Non-mammary neoplasias:
 - lipomas, mast cell tumours,
 - melanomas, subcut. hemangiosarcomas
- Histology needed for definitive diagnosis and prognosis



(Picture courtesy of Dr Juan Borrego DACVIM Oncology; cytology provided by IDEXX Laboratories)

(Simon et al. 2009; Sontas et al. 2012)

Surgery

- Surgical removal -> histology -> definitive diagnosis
 - Multiple masses ≠ not same diagnosis (66%) => remove them all = analyse them all
- General surgical approach guidelines:
 - Lumpectomy or mastectomy based on size <3 cm
 - Partial strip: >3cm / between mammary glands
 - Full strip only when too many lesions
- For masses >3 cm aim for 2 cm margins
 - (formaline fixed tissue \downarrow 30%)
- Mastectomy deep margins down to fascia
 - if recurrent /very infiltrative >>remove fascia too



(Picture courtesy of Juan Borrego, DACVIM Oncology)



Canine mammary tumours

Surgery: draining lymph node resection

• Double check lymphatic drainage anatomy prior to LN removal

Location of mammary Gland	Neoplastic Lymphatic Drainage
M1 - cranial thoracic	Axillary LN or sternal LN
M2 - caudal thoracic	Axillary LN or sternal LN
M3 - cranial abdominal	Axillary LN, superficial inguinal LN << medial iliac LN
M4 - caudal abdominal	Superficial inguinal LN << axillary LN
M5 - inguinal	Superficial inguinal LN < <popliteal ln<="" td=""></popliteal>

- Anatomical approach described for axillary LNs removal
 - Consider methilene blue to ease visualization
- If mastectomy of 5th or 4th+5th ... inguinal LN removal!



(Pictures courtesy of Dr. Manuel Jimenez DECVS)



Surgery

- Malignant mammary neoplasia based negative prognostic factors:
 - Wide margins (partial strip)+ lymph node (s) removal
 - No difference in survival for full strip / more postoperative complications
- When to biopsy:
 - Inflammatory mammary carcinoma suspected
 - Spread of localised mammary tumor to skin

(Horta et al. 2015, Evans et al. 2021)



Diagnosis: histology

- Is it needed? YES!!!
- 50% of canine mammary masses are benign
- For malignant mammary neoplasia:
 - Histologic grading = estimate of recurrence / metastasis risk
 - Some specific tumor types associated with malignant behaviour

(Sorenmo et al. 2020)



what's important in the histology report?

Grading & prognosis: Goldschmidt adapted WHO classification

- Applies to epithelial mammary neoplasia
- Grade I and II: risk of metastasis 19-30%, recurrence with narrow margins 9-12%
- Grade III: risk of metastasis 87%, recurrence 32%
- Lymphatic invasion present / absent
- Margin assessment incomplete /very narrow margins = recurrence risk
- Lymph node status

(Goldschmidt et al. 2011; Tran et al. 2016; Rasotto et al. 2017)



Diagnosis: histology

- Specific tumor types associated to agressive malignant behaviour: local and metastatic
- Comedocarcinoma, solid/anaplastic carcinoma
- Carcinosarcoma, osteosarcoma
- Inflammatory mammary carcinoma*
 - Diffuse skin lymphatic extension
 - Surgery contraindicated





Canine mammary tumours

Risk of metastasis: what to do?

- Target organs are lungs and draining lymph nodes (LNs)
- Action plan depends on estimated risk (tumor size, grade, stage)
- Periodic staging always for high met. risk / already present
- Staging:
 - Diagnostic imaging lungs
 - Locating draining LN(s) (might require imaging):
 - **REMOVAL** when high risk of metastasis after staging
 - If monitoring, initial FNA/size recording



Risk of metastasis: Chemotherapy, Radiotherapy

- Weak evidence of survival improvement in metastatic mammary neoplasia or inflammatory carcinomas
- No documented % decrease in metastatic rate compared to sx alone
 - Lack of optimally designed studies
- Doxorubicin, mitoxantrone, carboplatin, toceranib, COX-2 inhibitors,...
 - If used after complete wide resection /removal of regional LNs

Radiotherapy:

- Inflammatory mammary carcinoma
- Extensive skin involvement/nonresectable carcinomas
 - Improves survival compared to medical therapy alone



Benefits of spaying in dogs after diagnosis of mammary tumors?

- 50% of intact bitches will develop at least one mammary tumor
- Mammary tumors in spayed bitches are more frequently malignant
- Spaying at diagnosis of benign tumor will decrease by 50% risk of further mammary tumors
- OVHE does not change survival after diagnosis of malignant neoplasia
 - Exception adenocarcinomas grade II or high estradiol preoperative levels

(Sorenmo et al. 2000; Kristiansen et a. 2013; Kristiansen et al. 2016; Gedon et al. 2022)



References

Goldschmidt M, Peña L, Rasotto R, Zappulli V. Classification and grading of canine mammary tumors. Vet Pathol. 2011 Jan;48(1):117-31.

Rasotto R, Berlato D, Goldschmidt MH, Zappulli V. Prognostic Significance of Canine Mammary Tumor Histologic Subtypes: An Observational Cohort Study of 229 Cases. Vet Pathol. 2017 Jul;54(4):571-578.

Mainenti M, Rasotto R, Carnier P, Zappulli V. Oestrogen and progesterone receptor expression in subtypes of canine mammary tumours in intact and ovariectomised dogs. Vet J. 2014 Oct;202(1):62-8.

Gedon J, Wehrend A, Failing K, Kessler M. Canine mammary tumours: Size matters-a progression from low to highly malignant subtypes. Vet Comp Oncol. 2021 Dec;19(4):707-713.

Gedon J, Wehrend A, Kessler M. Ovariectomy reduces the risk of tumour development and influences the histologic continuum in canine mammary tumours. Vet Comp Oncol. 2022 Jun;20(2):476-483.

Evans BJ, Holt DE, Stefanovski D, Sorenmo KU. Factors influencing complications following mastectomy procedures in dogs with mammary gland tumors: 140 cases (2009-2015). J Am Vet Med Assoc. 2021 Feb 1;258(3):295-302.

Sorenmo KU, Shofer FS, Goldschmidt MH. Effect of spaying and timing of spaying on survival of dogs with mammary carcinoma. J Vet Intern Med. 2000 May-Jun;14(3):266-70.

Litterine-Kaufman J, Casale SA, Mouser PJ. Prevalence of malignancy in masses from the mammary gland region of dogs with single or multiple masses. J Am Vet Med Assoc. 2019 Oct 1;255(7):817-820.

Gunnes G, Borge KS, Lingaas F. A statistical assessment of the biological relationship between simultaneous canine mammary tumours. Vet Comp Oncol. 2017 Jun;15(2):355-365.

Peña L, Perez-Alenza MD, Rodriguez-Bertos A, Nieto A. Canine inflammatory mammary carcinoma: histopathology, immunohistochemistry and clinical implications of 21 cases. Breast Cancer Res Treat. 2003 Mar;78(2):141-8.

Langenbach A, Anderson MA, Dambach DM, Sorenmo KU, Shofer FD. Extraskeletal osteosarcomas in dogs: a retrospective study of 169 cases (1986-1996). J Am Anim Hosp Assoc. 1998 Mar-Apr;34(2):113-20.

Sorenmo KU, Rasotto R, Zappulli V, Goldschmidt MH. Development, anatomy, histology, lymphatic drainage, clinical features, and cell differentiation markers of canine mammary gland neoplasms. Vet Pathol. 2011 Jan;48(1):85-97.



References

Kristiansen VM, Nødtvedt A, Breen AM, Langeland M, Teige J, Goldschmidt M, Jonasdottir TJ, Grotmol T, Sørenmo K. Effect of ovariohysterectomy at the time of tumor removal in dogs with benign mammary tumors and hyperplastic lesions: a randomized controlled clinical trial. J Vet Intern Med. 2013 Jul-Aug;27(4):935-42.

Kristiansen VM, Peña L, Díez Córdova L, Illera JC, Skjerve E, Breen AM, Cofone MA, Langeland M, Teige J, Goldschmidt M, Sørenmo KU. Effect of Ovariohysterectomy at the Time of Tumor Removal in Dogs with Mammary Carcinomas: A Randomized Controlled Trial. J Vet Intern Med. 2016 Jan-Feb;30(1):230-41.

Tran CM, Moore AS, Frimberger AE. Surgical treatment of mammary carcinomas in dogs with or without postoperative chemotherapy. Vet Comp Oncol. 2016 Sep;14(3):252-62.

Arenas C, Peña L, Granados-Soler JL, Pérez-Alenza MD. Adjuvant therapy for highly malignant canine mammary tumours: Cox-2 inhibitor versus chemotherapy: a case-control prospective study. Vet Rec. 2016 Jul 30;179(5):125.

Rossi F, Sabattini S, Vascellari M, Marconato L. The impact of toceranib, piroxicam and thalidomide with or without hypofractionated radiation therapy on clinical outcome in dogs with inflammatory mammary carcinoma. Vet Comp Oncol. 2018 Dec;16(4):497-504.

Sorenmo K.U., Worley D.R., Zapulli V. Tumors of the mammary gland. In: Withrow & MacEwen's small animal clinical oncology. Sixth edition. St. Louis, Missouri: Elsevier, 2020.

Sontas BH, Yüzbaşıoğlu Öztürk G, Toydemir TF, Arun SS, Ekici H. Fine-needle aspiration biopsy of canine mammary gland tumours: a comparison between cytology and histopathology. Reprod Domest Anim. 2012 Feb;47(1):125-30.

Simon D, Schoenrock D, Nolte I, Baumgärtner W, Barron R, Mischke R. Cytologic examination of fine-needle aspirates from mammary gland tumors in the dog: diagnostic accuracy with comparison to histopathology and association with postoperative outcome. Vet Clin Pathol. 2009 Dec;38(4):521-8.

Horta RS, Figueiredo MS, Lavalle GE, Costa MP, Cunha RM, Araújo RB. Surgical stress and postoperative complications related to regional and radical mastectomy in dogs. Acta Vet Scand. 2015 Jun 24;57(1):34.

de Souza MCC, Flecher MC, Arrais FM, de Sena BV, Giuliano A, Horta RDS. Comparison of surgical resection of Axillary Lymph Nodes in Dogs with Mammary Gland Tumors with or without sentinel lymph node visualization with patent blue dye. Front Vet Sci. 2023 May 12;10:1149315.



Questions?

