

Hepatosplenic and Hepatocytotropic T-Cell Lymphoma: Two Distinct Types of T-Cell Lymphoma in Dogs

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Abstract

The clinical, clinicopathologic, and pathological findings of 9 dogs with T-cell lymphoma that involved the liver in the absence of peripheral lymphadenopathy were assessed. Seven dogs had hepatosplenic T-cell lymphoma (HS-TCL). Dogs with HS-TCL presented with hepato- and/or splenomegaly, regenerative anemia, thrombocytopenia, and hypoproteinemia. The clinical course was rapidly progressive with all dogs but 1 dead within 24 days of initial presentation. Neoplastic lymphocytes were centered on hepatic and splenic sinusoids and had a CD3+ (5/7), TCR α β+ (5/5), TCR γ δ+ (3/5), CD11d+ (6/7), granzyme B+ (5/7) immunophenotype. Bone marrow and lungs were consistently but variably involved. These findings closely resemble the human disease and support the classification of HS-TCL as a distinct World Health Organization entity in dogs. The remaining 2 dogs markedly differed in the pattern of hepatic involvement by neoplastic lymphocytes, which were not confined to hepatic sinusoids but invaded hepatic cords. In addition, neoplastic cells had a CD11d- immunophenotype, and clinicopathologic data indicated marked cholestasis and mild to absent anemia. Based on the distinct tropism of neoplastic lymphocytes for hepatocytes, the name hepatocytotropic T-cell lymphoma (HC-TCL) is proposed. Given the histomorphologic, clinicopathologic, and immunophenotypic differences, HC-TCL likely represents a separate biological entity rather than a histomorphologic variant of HS-TCL.

Keywords

emperipolesis, gamma delta T cell, hepatosplenic T cell lymphoma, hepatocytotropic T cell lymphoma, molecular clonality, WHO classification, PTCL-NOS

Considerable effort has been directed toward a generally accepted classification scheme for canine lymphomas.^{1,2} An appropriate classification scheme should adequately group tumors into clinicopathologic entities and serve as a basis for prediction of biological behavior and choice of treatment. With advances in research and diagnostic methods, classification schemes are subject to constant refinement and evolution.

The current most commonly accepted classification scheme for human lymphomas is the World Health Organization (WHO) system.¹⁸ The majority of WHO entities are defined by clinical, morphological, immunophenotypic, and genetic criteria. One such example is hepatosplenic T-cell lymphoma (HS-TCL). HS-TCL is defined as a T-cell lymphoma that is centered on the liver and spleen without significant involvement of peripheral lymph nodes.¹⁸ HS-TCL is thought to arise from splenic cytotoxic T cells and is predominantly of γ δ T-cell origin.^{1,4,19,21} Patients clinically present with splenomegaly and/or hepatomegaly, thrombocytopenia, and anemia. Histologically, neoplastic cells are predominantly found within splenic and hepatic sinusoids with concurrent bone marrow involvement in most cases.

HS-TCL has recently been adopted as a new entity in the canine WHO classification. However, so far only 2 cases have

been reported in dogs.^{18,19} Although these cases resembled the human disease, their singularity precluded identification of general characteristics of HS-TCL in dogs. Additional data are needed to more precisely characterize this tumor type and to validate its status as a distinct WHO entity in dogs.

In contrast to HS-TCL, other WHO entities are more heterogeneous and include a range of different tumor types that have not been further characterized. One such example, in humans as well as in dogs, is peripheral T-cell lymphoma—not otherwise specified (PTCL-NOS).^{18,19} In humans, this entity is



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The Lymphoid Neoplasms 3ed Ian Magrath,2010-07-30 This comprehensive and authoritative reference covers all aspects of the group of disorders collectively known as the lymphoid neoplasms The reader is taken through a description of its normal cellular origins and the molecular genetic abnormalities that can lead to this group of conditions a section of the book that has been considerably strengt

MRI and CT of the Female Pelvis Bernd Hamm,Rosemarie Forstner,2007-01-19 MRI and CT exquisitely depict the anatomy of the female pelvis and offer fascinating diagnostic possibilities in women with pelvic disorders This volume provides a comprehensive account of the use of these cross sectional imaging techniques to identify and characterize developmental anomalies and acquired diseases of the female genital tract Both benign and malignant diseases are considered in depth and detailed attention is also paid to normal anatomical findings and variants Further individual chapters focus on the patient with pelvic pain and the use of MRI for pelvimetry during pregnancy and the evaluation of fertility Throughout emphasis is placed on the most recent diagnostic and technical advances and the text is complemented by many detailed and informative illustrations All of the authors are acknowledged experts in diagnostic imaging of the female pelvis and the volume will prove an invaluable aid to everyone with an interest in this field **Morson and Dawson's Gastrointestinal Pathology** David W. Day,Jeremy R. Jass,Ashley B. Price,Neal A. Shepherd,James M. Sloan,Nicholas J. Talbot,Geraint T. Williams,Bryan F. Warren,2008-04-15 Morson and Dawson s Gastrointestinal Pathology is one of the Gold Standards of pathology textbooks It has been completely revised to incorporate the latest advances in this rapidly evolving field including the developments in gastric cancer and Helicobacter pylori and the revised classification of other common gastrointestinal conditions This new edition features a wealth of new material presented in full colour for the first time

Clinical Bone Marrow and Blood Stem Cell Transplantation Kerry Atkinson,2004 Drs Richard Champlin Jerome Ritz Willem Fibbe Per Ljungman and Malcom K Brenner join Kerry Atkinson as editors of this definitive reference on the clinical practice and underlying science of hematopoietic stem cell transplantation This third edition text is significantly revised and updated with 124 chapters balancing scientific explanations with practical information on patient care for all aspects of autologous syngeneic and allogeneic transplantation This edition includes 18 new chapters on significant topics such as plasticity of stem cells embryonic stem cells and nonmyeloablative conditioning regimens Thoroughly referenced through 2003 the chapters are divided into 15 sections including biological background and practical procedures clinical results transplant related and organ specific complications laboratory aspects and developing areas with a final breaking news chapter from this rapidly evolving field Over 170 internationally recognized experts contributed to this authoritative and practical text that is an essential resource for hematologists oncologists and transplant specialists

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Current Opinion in Oncology, 1998 Herpesviruses and Immunity Peter G. Medveczky, Herman Friedman, Mauro Bendinelli, 2006-04-11 Although virology and immunology are now considered separate disciplines history shows that these areas of investigation always overlapped and one cannot really exist without the other This trend has become particularly significant and fruitful in the past few years in the area of herpesvirus research The genomes of the most important herpesviruses have been sequenced a significant portion of their genes have been identified and many secrets of regulation of gene expression have been unraveled Now this progress sets the stage for a true revolution in herpesvirus research analysis of interactions between the host and the virus Because herpesviruses can induce suppress and fool the immune system the most productive herpesvirologists are also expert immunologists and the current results of this interdisciplinary effort are truly remarkable Because herpesviruses cause many important human diseases the development of vaccines against these agents is a very significant goal This effort is also very challenging because of the complexity of herpesviruses and the lack of sufficient information about immune responses The remarkable ability of herpesviruses to escape immune responses is another feature that brings immunology and virology together Herpesviruses code many proteins that interact with and down regulate some key elements of the immune system This property of herpesviruses represents a major challenge in developing strategies against these viruses On the positive side these viral proteins also provide novel tools for analyzing specific immune reactions and molecular mechanisms

Clinical Nuclear Medicine Hans-Jürgen Biersack, Leonard M. Freeman, 2008-01-03 The modern era of radionuclide imaging and therapy is well into its seventh decade During this era many national and international textbooks have been published in an attempt to educate not only the practitioners of our medical discipline but also referring physicians and medical students Some of the more recent large multiauthored texts such as those by Ell and Ghambir Sandler et al and Henkin et al provide us with very comprehensive reference sources while some of the smaller texts totally written by two or three individuals e.g. Mettler Guibert and Ziessman O'Malley Thrall have achieved popularity with radiology residents and other physicians in training The concept of Clinical Nuclear Medicine arose 3 years ago from a conversation between the editors who have been close friends for many years We have always felt that our relationship epitomizes one of the major strengths of nuclear medicine which is the very close ties and spirit of educational cooperation that exist between international colleagues We all share the same aim of doing whatever we can to optimize patient care whether it be by introducing new pharmaceuticals and instruments or by developing new techniques or approaches to performing our broad spectrum of clinical procedures Nuclear medicine

physicians have almost uniformly been willing to share their expertise at national and international meetings The international nuclear medicine community unlike many other larger specialty areas has remained relatively small It was within this spirit that Clinical Nuclear Medicine was born

The Lymphomas George Peter Canellos, Thomas Andrew Lister, Bryan D. Young, 2006 Edited and written by world authorities this text encompasses all aspects of Hodgkin s disease non Hodgkin s lymphomas and related disorders Designed to be the Bible for both clinicians and scientists who deal with lymphoma the book is packed with useful summary tables and abundant illustrations and translates cutting edge discoveries into state of the art management guidelines The preeminent names in research and practice bring together today s entire spectrum of knowledge from epidemiology classification and prognosis through diagnosis and therapy including the latest scientific advances such as the impact of molecular biologic and genetic discoveries and new biologic markers Covers molecular biology bone marrow transplantation lymphoma in AIDS and Hodgkins disease Provides in depth coverage of the underlying biology of the disease as well as etiology pathology diagnosis and treatment Discusses advances in the treatment of diseases including radiotherapy radiolabeled monoclonal antibodies treatment of cutaneous lymphoma and metastatic disease Contains precise clinical how tos for managing bone marrow transplantation and diseases that resemble lymphomas Includes the recently codified new WHO classification of all lymphoid entities Offers new and updated information on the biology and recategorization morphologic clinical genotypic immunophenotypic of distinct lymphoma entities

The Journal of Immunology, 2005

Molecular Biology of B Cells Tasuku Honjo, Michael Reth, Andreas Radbruch, Frederick Alt, Michael Neuberger, 2004-02-19 Molecular Biology of B Cells is a comprehensive reference to how B cells are generated selected activated and engaged in antibody production All these developmental and stimulatory processes are described in molecular and genetic terms to give a clear understanding of complex phenotypes The molecular basis of many diseases due to B cell abnormality is also discussed This definitive reference is directed at research level immunologists molecular biologists and geneticists

The Cancer Handbook Malcolm Alison, 2007

Archives of Pathology & Laboratory Medicine, 2002

Tumors of the Intestines Robert H. Riddell, 2003

Radiation Oncology James Daniel Cox, K. Kian Ang, 2003 The New Edition of this highly respected resource equips readers with the very latest clinical guidelines in radiation oncology 52 leading experts many of them new to this edition present their preferred management approaches for all body regions emphasizing how to deliver precise targeted radiation doses while sparing healthy tissue

Cancer Research, 2005-06

Uveitis Robert B. Nussenblatt, Scott M. Whitcup, 2004 The New Edition of this practical reference has been completely revised and updated to deliver cutting edge guidance on the latest medical pharmacological and surgical treatments of uveitis Known for its balanced scientific and clinical approach it comprehensively covers diagnostic testing differential diagnosis treatment immunology epidemiology clinical manifestations and therapeutic approaches Covers the medical pharmacological and surgical treatments of uveitis as viewed by two world class clinicians Provides a wealth of vividly illustrated case studies

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Williams Hematology Malignant Lymphoid Diseases Oliver W Press, Marshall A. Lichtman, John P. Leonard, 2018-01-12 Leukemias and lymphoid diseases as only Williams can cover them Featuring content derived from Williams Hematology Ninth Edition this concise full color resource delivers comprehensive and current coverage of leukemias and lymphoid diseases More than a sectional reprint the book includes updated science and treatment recommendations not found in the 2015 release of Williams Hematology Ninth Edition Perfect for use at the point of care Williams Malignant Lymphoid Diseases maintains a strong clinical focus with detailed discussion of clinical presentation patient evaluation and diagnosis genetic basis therapy and management A handy quick summary appears at the beginning of each chapter Includes the latest advances in common treatments of leukemias such as targeted therapy immunotherapy bone marrow hematopoietic stem cell transplantation and monoclonal antibodies

The MD Anderson Manual of Medical Oncology, Second Edition Hagop M. Kantarjian, Robert A. Wolff, 2011-04-22 State of the art approaches to cancer management from the world's 1 oncology center The perfect clinical desk manual from the experts at MD Anderson Cancer Center Read the outstanding reviews of the previous edition As a detailed guide the book offers practitioners a clear user friendly approach to the care of patients with cancer New England Journal of Medicine The portions of the text that summarize the chemotherapeutic literature are helpful and potentially useful to a busy medical oncologist The graphic representations and flow charts are of high quality are well designed and easy to follow and are clinically useful A number of good photographs have been provided that illustrate pathologic conditions the Manual is well written has extremely useful graphics and represents a useful entry point for those interested in the management of malignant diseases International Journal of Radiation Oncology Biology Physics MD Anderson Manual of Medical Oncology fulfills its goals of helping clinicians quickly evaluate therapeutic regimens and possibilities Surgeons and radiation oncologists will find the text useful as a reference guide to what the medical oncologist can do and how the medical oncologist views their disciplines The text is also of considerable help for physicians studying for board certification or recertification in medical oncology Practicing medical oncologists would do well to have it in their personal libraries Journal of the American Medical Association MD Anderson Manual of Medical Oncology offers clinicians an authoritative easy to follow guide to the medical management of patients with cancer and its complications Featuring concise but practical text and a large number of decision making algorithms the book gives physicians a current templated approach to each disease and situation backed by the clinical expertise and authority of world recognized oncology researchers practitioners Key Features REVISED completely updated coverage of the quickly evolving world of cancer therapeutics NEW chapters include Expert Commentary Sidebars that provide clinical insights from leading subspecialists in radiation therapy surgical oncology

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