

Return of Organization Exempt From Income Tax

Under section 501(c), 527, or 4947(a)(1) of the Internal Revenue Code (except black lung benefit trust or private foundation)

2003
Open to Public

The organization may have to use a copy of this return to satisfy state reporting requirements.

merr	ar never	de Service Missing in the Country of		mspection
A F	or the 2	2003 calendar year, or tax year beginning and ending		
В	heck if	I Flease I	mployer ic	dentification number
		Juse HS AMERICAN KENNEL CLUB CANINE		
X	Addres	print or HEALTH FOUNDATION, INC.	13-38	813813
<u>_</u>]Name Jchang∈ ∏Initial) See)	elephone	
<u>_</u>	_ireturn	Specific 5580 CENTERVIEW DRIVE		<u>682-9696</u>
<u>_</u>	Final return Amend	\int tions City or town, state or country, and $ZIP + 4$	Accounting met	
<u>_</u>	⊒return	RALEIGH, NC 27000-3390	Other (specify)	
<u></u>	Applica	must attach a completed Schedule & (Form 90) or 990-E7)		<u>~</u>
		H(a) is this a group return		
		:▶WWW.AKCCHF.ORG H(b) If "Yes," enter numb		
		ation type (check only one) ► X 501(c) (3) ◀ (insert no) 4947(a)(1) or 527 H(c) Are all affiliates including the second of th		N/A Yes No
		ere \triangleright in the organization's gross receipts are normally not more than \$25,000. The $ $ H(d) is this a separate re-	turn filed b	y an or-
		tion need not file a return with the IRS; but if the organization received a Form 990 Package ganization covered		ruling? Yes X No
	tne m	ail, it should file a return without financial data. Some states require a complete return. [Group Exemption N		
				tion is not required to attach
		ceipts: Add lines 6b, 8b, 9b, and 10b to line 12 ★ 4, 250, 563 Sch. B (Form 990, 9	90-62, 01	990-PF).
PE	rt I	Revenue, Expenses, and Changes in Net Assets or Fund Balances		
	1	Contributions, gifts, grants, and similar amounts received:	. { }	
	a	Direct public support 1a 2,570,338	∸ }	
	þ	Indirect public support 1b	-	
	C	Government contributions (grants)	\dashv \downarrow	0 570 220
	d	Total (add lines 1a through 1c) (cash \$ 2,483,457. noncash \$ 86,881.)	1d	2,570,338.
	2	Program service revenue including government fees and contracts (from Part VII, line 93)	2	
	3	Membership dues and assessments	3	
	4	Interest on savings and temporary cash investments	5	100,341.
	5	Dividends and interest from securities	3	100,341.
	6 a	Gross rents Less: rental expenses 6a 6b	-	
	b	Cutation and Control of the Control	ا و ا	
	, c	Net rental income or (loss) (subtract line 6b from line 6a)	6c 7	
ř	7	Other investment income (describe Gross amount from sales of assets other (A) Securities (B) Other	+	
Revenue	Oa	4 42 5 444	- I	
Be		than inventory		
	b	Gain or (loss) (attach schedule) 482,322.>8c <19,503		
	d	CITATION OF THE CITATION OF TH	80	<101,825.>
	9	Special events and activities (attach schedule). If any amount is from gaming, check here	100	<u> </u>
		Gross revenue (not including \$ 133,116. of contributions	\. · · · \	
	•	reported on line 1a) 9a 60,900		
	ь	Less; direct expenses other than fundraising expenses	7	
	c	Net income or (loss) from special events (subtract line 9b from line 9a) SEE STATEMENT 3	9c	60,900.
	10 a	Gross sales of inventory, less returns and allowances		
	ь	Lace; cost of goods sold	7 '	
	C	Gross prefit of (loss line) take of inventory (attach schedule) (subtract line 10b from line 10a)	10c	
	11	Other revenue Grown Pane Vit, line 103) 2	11	19,539.
,	12	Total revenue (add lines 1d, 2,384, 5), 66, 7, 8d, 9c, 10c, and 11)	12	2,649,293.
)	13	Proprant services from the 44, column (H)	13	1,981,919.
Expenses	14	Managament and general (from line 44, column (C))	14	535,098.
Sense	15	Fundaising (from upa 44 Galumn (D))	15	232,844.
ă	16	Payments to affiliates (attach schedule)	16	
	17	Total expenses (add lines 16 and 44, column (A))	17	2,749,861.
	18	Excess or (deficit) for the year (subtract line 17 from line 12)	18	<100,568.>
ets	19	Net assets or fund balances at beginning of year (from line 73, column (A))	19	2,300,394.
Net Assets	20	Other changes in net assets or fund balances (attach explanation) SEE STATEMENT 4	20	421,907.
	21	Net assets or fund balances at end of year (combine lines 18, 19, and 20)	21	2,621,733.
3230 12-1	01 7-03	LHA For Paperwork Reduction Act Notice, see the separate instructions.		Form 990 (2003) (1

P	Statement of All org	anıza	ations must complete columnianizations and section 494	nn (A) Columns (B), (C), and 7(a)(1) nonexempt charitable	i (D) are required for section	501(c)(3) Page 2
	Do not include amounts reported on line) org	(A) Total	(B) Program	(C) Management	(D) Fundraising
	6b, 8b, 9b, 10b, or 16 of Part I	<u> </u>	(A) Total	services	` and general	(b) I dilutaising
22	Grants and allocations (attach schedule) cash \$1570391. noncash \$	22	1,570,391.	1,570,391.		
22	Specific assistance to individuals (attach schedule)	23	1,3/0,391.	1,3/0,3910		
	Benefits paid to or for members (attach schedule)	24				
25		25	143,461.	75,981.	50,279.	17,201.
	Other salaries and wages	26	152,904.		39,321.	30,560.
	Pension plan contributions	27	8,918.	1	2,624.	1,495.
	Other employee benefits	28	149,521.	1	83,591.	19,564.
	Payroll taxes	29	30,641.	16,316.	9,291.	5,034.
	Professional fundraising fees	30			•	
31		31				,
32	Legal fees	32				
	Supplies	33	4,509.	1,353.	2,180.	976.
34	Telephone	34	6,889.	2,597.	2,906.	1,386.
35	Postage and shipping	35	9,764.	1,793.	5,036.	2,935.
36	Occupancy	36	29,674.	10,682.	14,245.	4,747.
37	Equipment rental and maintenance	37	21,325.	4,820.	12,606.	3,899.
38	Printing and publications	38	47,311.	28,974.	9,213.	9,124.
39	Travel	39	29,642.	9,167.	9,287.	11,188.
40	Conferences, conventions, and meetings	40	167,075.	71,069.	21,565.	74,441.
41	Interest	41				
42	Depreciation, depletion, etc. (attach schedule)	42	19,346.		15,345.	4,001.
43	Other expenses not covered above (itemize):					
	a	43a				
	b	43b				
	c	43c				
	d	43d				
	e SEE STATEMENT 5	43e			257,609.	46,293.
44	Total functional expenses (add lines 22 through 43), Organizations completing columns (B)-(D), carry these totals to lines 13-15 int Costs. Check		2,749,861.	1,981,919.	535,098.	232,844.
lf " (iii	e any joint costs from a combined educational campai Yes," enter (i) the aggregate amount of these joint cos i) the amount allocated to Management and general \$ Part III Statement of Program Servi	sts \$; and	(ii) the amount allocated to (iv) the amount allocated to	Program services \$	Yes X No ,
	nat is the organization's primary exempt purpose?					
**	iat is the organization a primary exempt purpose.		BE SIKIEMENI			Program Service
	organizations must describe their exempt purpose achievement					Expenses (Required for 501(c)(3) and
	nievements that are not measurable (Section 501(c)(3) and (4) or ecations to others)	ganız	ations and 4947(a)(1) nonexempt	charitable trusts must also enter	the amount of grants and	(4) orgs, and 4947(a)(1) trusts, but optional for others)
а	SEE STATEMENT 7					
				Grants and allocations \$	1,570,391.)	1,981,919.
b			<u> </u>			
				Grants and allocations \$)	
C						
_				Grants and allocations \$)	
C						
				(Grants and allocations \$	· · · · · · · · · · · · · · · · · · ·	
-	Other program services (attach schedule)			Grants and allocations \$	1	
				<u>'</u>		1 001 010
f	Total of Program Service Expenses (should equal	line 4	14, column (B). Program se	rvices)	•	1,981,919.

Form 990 (2093)

Part IV Balance Sheets

		e required, attached schedules and amour ld be for end-of-year amounts only	ts within the description column	(A) Beginning of year	(B) End of year
	45	Cash - non-interest-bearing		45	282,175.
S	46	Savings and temporary cash investments		966,590. 46	
	47 a	Accounts receivable	47a		
Ì	b	Less: allowance for doubtful accounts	47b	47	C
	48 a	Pledges receivable	48a 357,653.		
ŀ	то a h	Less: allowance for doubtful accounts	48b	458,776. 48	357,653
	49	Grants receivable		49	
	50	Receivables from officers, directors, trustees,			
		and key employees		50)
3	51 a	Other notes and loans receivable	51a		
{ (b	Less: allowance for doubtful accounts	51b	51	C
	52	Inventories for sale or use	_	52	
	53	Prepaid expenses and deferred charges		6,744. 53	
	54		MT 9 ► Cost X FMV	2,968,443. 54	3,128,407
	55 a				
		equipment basis	55a		
		Less: accumulated depreciation	55b	55	
ļ	56	Investments - other	SEE STATEMENT 10	0 . 56	424,472
1	57 a	Land, buildings, and equipment basis	57a 92,171.	44 177 5	76 700
	58	Less: accumulated depreciation Other assets (describe ▶	57b 15,381. SEE STATEMENT 11)	44,177. 57 95,598. 58	
	30	Other assets (describe	SEE STATEMENT II	75,550. 38	00,711
	59	Total assets (add lines 45 through 58) (must e	qual line 74)	4,540,328. 59	5,163,872
	60	Accounts payable and accrued expenses		111,090.60	
	61	Grants payable		2,095,744. 61	1 2,446,818
	62	Deferred revenue		62	2
Liabilities	63	Loans from officers, directors, trustees, and ke	y employees	63	3
	64 a	Tax-exempt bond liabilities		64	
<u> </u>	t	Mortgages and other notes payable		64	
	65	Other liabilities (describe)	33,100. 65	5
	66	Total liabilities (add lines 60 through 65)		2,239,934. 66	2,542,139
ヿ		nizations that follow SFAS 117, check here	X and complete lines 67 through		
.		69 and lines 73 and 74.			
ဗိ	67	Unrestricted		<773,059.>67	
9	68	Temporarily restricted		2,023,033. 68	
Š	69	Permanently restricted		1,050,420.69	1,051,441
5	Orga	nizations that do not follow SFAS 117, check he	ere and complete lines		
L X		70 through 74.			
2	70	Capital stock, trust principal, or current funds	-	70	
200	71	Paid-in or capital surplus, or land, building, and		7	
∢ ∣	72	Retained earnings, endowment, accumulated in		72	2
ای	70	Total net assets or fund balances (add lines 6	7 through 69 or lines 70 through 72;		
Net Assets or Fund Balances	73	column (A) must equal line 19, column (B) mu		2,300,394. 73	2,621,733

Form 990 is available for public inspection and, for some people, serves as the primary or sole source of information about a particular organization. How the public perceives an organization in such cases may be determined by the information presented on its return. Therefore, please make sure the return is complete and accurate and fully describes, in Part III, the organization's programs and accomplishments.

AMERICAN KENNEL CLUB CANINE

Form 990 (2003) HEALTH FOUN							<u> 3813 </u>	Page 4
Part IV-A Reconciliation of Revenue por Financial Statements with Research	er Audited evenue per	Part	Financia	iliation of Exp al Statements				
Return a Total revenue, gains, and other support per audited financial statements b Amounts included on line a but not on line 12, Form 990 (1) Net unrealized gains on investments \$ 421,907. (2) Donated services and use of facilities \$ 42,075. (3) Recoveries of prior year grants \$ (4) Other (specify): STMT 12 \$ 70,000. Add amounts on lines (1) through (4) b c Line a minus line b c d Amounts included on line 12, Form 990 but not on line a: (1) Investment expenses	533,982. 2,670,130.	(2) (3) (4) <u>S'</u> c	Return Total expenses and lo audited financial state. Amounts included on line 17, Form 990. Donated services and use of facilities. Prior year adjustment: reported on line 20, Form 990. Losses reported on line 20, Form 990. Other (specify): TMT 13. Add amounts on lines. Line a minus line b. Amounts included on 990 but not on line a: Investment expenses.	### ##################################		b	132,5 2,749,8	912.
not included on line 6b, Form 990 \$		(2)	not included on line 6b, Form 990 Other (specify):	\$ \$				
Add amounts on lines (1) and (2)	<20,837.	>	Add amounts on lines		>	d		0.
e Total revenue per line 12, Form 990	2 640 202	е	Total expenses per lin (line c plus line d)	e 17, Form 990		.	2740 0	0.6.1
(line c plus line d) Part V List of Officers, Directors, Trus	2,649,293. tees, and Key E		<u> </u>	e even if not comper	nsated)	e	2,749,8	<u>, , , , , , , , , , , , , , , , , , , </u>
		(B) Tif	le and average hours	(C) Compensation		tribution	s to (E) Exp	ense
(A) Name and address		pe	r week devoted to position	(If not paid, enter	plans	& deferre	accour	it and wances
SEE STATEMENT 15				143,461.	39	,60	2. 33,9	952.
								
						_		
		-			-			
							-	
· · · · · · · · · · · · · · · · · · ·	·	-			 			
75 Did any officer, director, trustee, or key employee receive	e aggregate compensati	on of m	ore than \$100,000 fro	m your organization	and all	related		
organizations, of which more than \$10,000 was provided					X No			
323031 12-17-03							Form 990	(2003)

AMERICAN KENNEL CLUB CANINE

Form	990 (2003) HEALTH FOUNDATION, INC. 13-3813	813		Page 5
Par	t VI Other Information		Yes	No
76	Did the organization engage in any activity not previously reported to the IRS? If "Yes," attach a detailed description of each activity	76		X
77	Were any changes made in the organizing or governing documents but not reported to the IRS?	77	Х	
	If "Yes," attach a conformed copy of the changes			
78 a	Did the organization have unrelated business gross income of \$1,000 or more during the year covered by this return?	78a		X
b	If "Yes," has it filed a tax return on Form 990-T for this year? N/A	78b		
79	Was there a liquidation, dissolution, termination, or substantial contraction during the year?	79		X
	If "Yes," attach a statement			
80 a	Is the organization related (other than by association with a statewide or nationwide organization) through common membership,			1
	governing bodies, trustees, officers, etc., to any other exempt or nonexempt organization?	80a		<u>X</u>
b	If "Yes," enter the name of the organization			
	and check whether it is exempt or nonexempt.			
81 a	Enter direct or indirect political expenditures. See line 81 instructions 81a 0.			
þ	Did the organization file Form 1120-POL for this year?	81b		X
82 a	Did the organization receive donated services or the use of materials, equipment, or facilities at no charge or at substantially less than			
	fair rental value?	82a	X	
b	If "Yes," you may indicate the value of these items here. Do not include this amount as revenue in Part I or as an			
	expense in Part II. (See instructions in Part III.)	4		
83 a	Did the organization comply with the public inspection requirements for returns and exemption applications?	83a	Х	
þ	Did the organization comply with the disclosure requirements relating to quid pro quo contributions?	83b	X	
84 a	Did the organization solicit any contributions or gifts that were not tax deductible? N/A	84a		
b	If "Yes," did the organization include with every solicitation an express statement that such contributions or gifts were not			
	tax deductible? N/A	84b		
85	501(c)(4), (5), or (6) organizations a Were substantially all dues nondeductible by members? N/A	85a	L	<u> </u>
b	Did the organization make only in-house lobbying expenditures of \$2,000 or less? N/A	85b		<u> </u>
	If "Yes" was answered to either 85a or 85b, do not complete 85c through 85h below unless the organization received a waiver for proxy tax			
	owed for the prior year.		~	
C	Dues, assessments, and similar amounts from members 85c N/A	4		
d	Section 162(e) lobbying and political expenditures 85d N/A	4		
е	Aggregate nondeductible amount of section 6033(e)(1)(A) dues notices 85e N/A	4		
f	Taxable amount of lobbying and political expenditures (line 85d less 85e) 85f N/A		\ \ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\	k.
g	Does the organization elect to pay the section 6033(e) tax on the amount on line 85f? N/A	85g	<u> </u>	
h	If section 6033(e)(1)(A) dues notices were sent, does the organization agree to add the amount on line 85f to its reasonable estimate of dues			
	allocable to nondeductible lobbying and political expenditures for the following tax year? N/A	85h	 	
86	501(c)(7) organizations Enter: a Initiation fees and capital contributions included on line 12 86a N/A	-		
D	Gross receipts, included on line 12, for public use of club facilities 86b N/A	┨	:	
87	501(c)(12) organizations. Enter: a Gross income from members or shareholders 87a N/A	-		
b	Gross income from other sources. (Do not net amounts due or paid to other sources			
	against amounts due or received from them.) 87b N/A	╣`	1	
88	At any time during the year, did the organization own a 50% or greater interest in a taxable corporation or partnership,			
	or an entity disregarded as separate from the organization under Regulations sections 301.7701-2 and 301.7701-37	00		v
90 -	If "Yes," complete Part IX 501(c)(3) organizations. Enter: Amount of tax imposed on the organization during the year under:	88		<u>X</u>
oa g	section 4911 O • ; section 4912 O • ; section 4955 O • ;			
ь	501(c)(3) and 501(c)(4) organizations Did the organization engage in any section 4958 excess benefit			j
U	transaction during the year or did it become aware of an excess benefit transaction from a prior year?			
	If "Yes," attach a statement explaining each transaction	89b		х
•	Enter: Amount of tax imposed on the organization managers or disqualified persons during the year under	030	1	
·	sections 4912, 4955, and 4958			0.
d	Enter: Amount of tax on line 89c, above, reimbursed by the organization			0.
	List the states with which a copy of this return is filed SEE STATEMENT 16			
	Number of employees employed in the pay period that includes March 12, 2003			8
91	The books are in care of ► AKC CANINE HEALTH FOUNDATION, INC. Telephone no. ► 888-68	32-9	696	
	<u> </u>			
	Located at ► 5580 CENTERVIEW DRIVE, RALEIGH, NC ZIP+4 ► 2	2760	6-3	390
92	Section 4947(a)(1) nonexempt chantable trusts filing Form 990 in lieu of Form 1041- Check here		▶[
	and enter the amount of tax-exempt interest received or accrued during the tax year	N/	Α	
32304 12-17-	1 03	For	n 990	(2003)

m 990 (2003) HEA	ALTH FOUNDA ne-Producing Ac			ctions)	13-3	813813 Page
ote: Enter gross amounts unless of			d business income		by section 512, 513, or 514	(5)
dicated 3 Program service revenue	nerwise	(A) Business code	(B) Amount	(C) Exclu- sion code	(D) Amount	(E) Related or exempt function income
ab						
b c	1					
d						
f Medicare/Medicaid payments						
g Fees and contracts from governmen	t agencies					
Membership dues and assessments						
Interest on savings and temporary c	ash investments					
Dividends and interest from securities	es			14	100,341.	
Net rental income or (loss) from real	estate:					
a debt-financed property	<u> </u>					
not debt-financed property	-					
Net rental income or (loss) from per-	sonal property					
Other investment income	-					
Gain or (loss) from sales of assets					101 005	
other than inventory	<u> </u>			18	<101,825.>	-
Net income or (loss) from special ev				01	60,900.	
Gross profit or (loss) from sales of II	nventory					······································
Other revenue:					10 546	
MISCELLANEOUS				01 15	10,546.	
ROYALTY INCOME				12	8,993.	
<u> </u>						
d						
Subtotal (add columns (B), (D), and	/E\\		0		78,955.	C
Total (add line 104, columns (B), (D)	-		Ų.	•11	10,233	78,955
e: Line 105 plus line 1d, Part I, sh		nt on line 12	Part I		_	10,55
art VIII Relationship of A	ctivities to the	Accompli	shment of Exem	pt Purpo	oses (See page 34 of the II	istructions.)
ne No. Explain how each activity for				d important	tly to the accomplishment of	the organization's
exempt purposes (other than	n by providing funds for	r such purpos	es).			
			<u></u>	······		
		Subsidiari		iea Enti	ties (See page 34 of the in	structions.)
(A) Name, address, and EIN of corporation partnership, or disregarded entity	(B) Percentage of ownership interest	t	(C) Nature of activities		(D) Total income	(E) End-of-year assets
	9/	6				,
N/A	9/	6				
	9/					
	9/					
art X Information Rega	rding Transfers	Associat	ed with Persona	l Benefi	t Contracts (See page	
i) Did the organization, during the yea	•	-			l benefit contract?	Yes X
) Did the organization, during the yea	ar, pay premiums, direc	tly or indirectly	y, on a personal benefit o	contract?		Yes X

Note: If "Yes" to (b), file Form 8870	and Form 4720	(see instructions)					
			mation of which pro	eparer has any knibwi	to the best of my knowle edge 2740 CBA	-	pelief, it is true, Preschout
		Davle		Type or print na	ime and title.		
				Date	Check if	Prepar	rer's SSN or PTIN

(Form 990 or 990-EZ)

Department of the Treasury

Internal Revenue Service Name of the organization **Organization Exempt Under Section 501(c)(3)**

(Except Private Foundation) and Section 501(e), 501(f), 501(k), 501(n), or Section 4947(a)(1) Nonexempt Charitable Trust

Supplementary Information-(See separate instructions.)

▶ MUST be completed by the above organizations and attached to their Form 990 or 990-EZ

AMERICAN KENNEL CLUB CANINE

OMB No 1545-0047

Employer identification number

		<u> 13 38138</u>	313
=	ficers, Directo	rs, and Trus	tees
(b) Title and average hours per week devoted to position	(c) Compensation	(d) Contributions to employee benefit plans & deferred compensation	(e) Expense account and othe allowances
DIR. GRANTS			
40	58,033.	10,177	4,974.
		al Services	
than \$50,000		service	(c) Compensation
		, ,	57,396.
		I	66 214
<u>11 48820 (</u> 	GIC PLANNI	NG	66,314.
			
	cr None.') (b) Title and average hours per week devoted to position DIR. GRANTS 40 Dendent Contractors or firms). If there are none, enter than \$50,000	oyees Other Than Officers, Director or 'None.') (b) Title and average hours per week devoted to position DIR. GRANTS 40 58,033. Oendent Contractors for Professions firms). If there are none, enter 'None.') than \$50,000 (b) Type of standard REF OH 44024 CONSULTANT	(b) Title and average hours per week devoted to position DIR. GRANTS 40 58,033. 10,177 0 Dendent Contractors for Professional Services or firms). If there are none, enter "None.") than \$50,000 ANNUAL REPORT, BROCHURES, ARTICL CONSULTANT/STRATE

AMERICAN KENNEL CLUB CANINE

1	e A (From 990 or 990-EZ) 2003 HEALTH FOUNDATION, INC. 13-38:			age 2
Part	Statements About Activities (See page 2 of the instructions.)	Y	'es	No
	ng the year, has the organization attempted to influence national, state, or local legislation, including any attempt to influence			
	lic opinion on a legislative matter or referendum? If "Yes," enter the total expenses paid or incurred in connection with the			
	sying activities 🕨 \$ (Must equal amounts on line 38, Part VI-A,		Ī	
	ne i of Part VI-B.)	1		<u>X</u>
_	anizations that made an election under section 501(h) by filing Form 5768 must complete Part VI-A. Other organizations checking			
	," must complete Part VI-B AND attach a statement giving a detailed description of the lobbying activities.			
	ing the year, has the organization, either directly or indirectly, engaged in any of the following acts with any substantial contributors,			
	tees, directors, officers, creators, key employees, or members of their families, or with any taxable organization with which any such			
•	son is affiliated as an officer, director, trustee, majority owner, or principal beneficiary? (If the answer to any question is "Yes,"			
	ich a detailed statement explaining the transactions) SEE STATEMENT 17			
a Sale	e, exchange, or leasing of property?	2a		X
b Len	ding of money or other extension of credit?	2b		Х
2 20.7				
c Furr	nishing of goods, services, or facilities?	2c		Х
d Pav	ment of compensation (or payment or reimbursement of expenses if more than \$1,000)? SEE PART V, FORM 990	2d	х	
uray	ment of compensation (or payment of reimbursement of expenses it more than \$4,000).	20	7	
e Tran	nsfer of any part of its income or assets?	2e		X
3 a Dov	you make grants for scholarships, fellowships, student loans, etc.? (If "Yes," attach an explanation of how			
you	determine that recipients qualify to receive payments.)	3a		X
b Doy	you have a section 403(b) annuity plan for your employees?	3b		X
	you maintain any separate account for participating donors where donors have the right to provide advice he use or distribution of funds?	4	х	
Part		1 7 1		
The orga	inization is not a private foundation because it is: (Please check only ONE applicable box.)	•		
5 [A church, convention of churches, or association of churches. Section 170(b)(1)(A)(i).			
6	A school. Section 170(b)(1)(A)(II). (Also complete Part V.)			
7	A hospital or a cooperative hospital service organization. Section 170(b)(1)(A)(iii).			
8	A Federal, state, or local government or governmental unit. Section 170(b)(1)(A)(v).			
9	A medical research organization operated in conjunction with a hospital. Section 170(b)(1)(A)(III). Enter the hospital's name, city,			
_	and state 🕨			
10	An organization operated for the benefit of a college or university owned or operated by a governmental unit. Section 170(b)(1)(A)(iv).		
_	(Also complete the Support Schedule in Part IV-A.)			
11a	An organization that normally receives a substantial part of its support from a governmental unit or from the general public.			
	Section 170(b)(1)(A)(vi). (Also complete the Support Schedule in Part IV-A.)			
11b	A community trust. Section 170(b)(1)(A)(vi). (Also complete the Support Schedule in Part IV-A.)			
12 L	An organization that normally receives: (1) more than 33 1/3% of its support from contributions, membership fees, and gross			
	receipts from activities related to its charitable, etc., functions - subject to certain exceptions, and (2) no more than 33 1/3% of			
	its support from gross investment income and unrelated business taxable income (less section 511 tax) from businesses acquired			
	by the organization after June 30, 1975. See section 509(a)(2). (Also complete the Support Schedule in Part IV-A.)			
13 [An organization that is not controlled by any disqualified persons (other than foundation managers) and supports organizations desc	rıbed ın:		
	(1) lines 5 through 12 above; or (2) section 501(c)(4), (5), or (6), if they meet the test of section 509(a)(2). (See section 509(a)(3).)			
	Provide the following information about the supported organizations. (See page 5 of the instructions.)			
	(a) Name(s) of supported organization(s)	(b) Line fron	numl n abo	
14	An organization organized and operated to test for public safety. Section 509(a)(4). (See page 6 of the instructions.)	 .		
٠٠ [

323111 12-05-03

f	Public support percentage (line	26e (numerator) divid	ded by line 26c (denominator))	▶ 261	46.1084%
27	Organizations described on line	12: a For amounts inc	cluded in lines 15, 16, and 17 that were received from a "disqualified pe	rson," pre	pare a list for your
	records to show the name of, an	d total amounts receive	ed in each year from, each "disqualified person." Do not file this list wit	h your ret	urn. Enter the sum of
	such amounts for each year:	N/A			
	(2002)	(2001)	(2000)	(1999)	

For any amount included in line 17 that was received from each person (other than "disqualified persons"), prepare a list for your records to show the name of, and amount received for each year, that was more than the larger of (1) the amount on line 25 for the year or (2) \$5,000. (Include in the list organizations described in lines 5 through 11, as well as individuals.) Do not file this list with your return. After computing the difference between the amount received and the larger amount described in (1) or (2), enter the sum of these differences (the excess amounts) for each year. (2002)(2001)(2000)(1999)

;	Add: Amounts from column (e) for lines:	15	16					
	17	20	21		•	27c	N/A	
1	Add: Line 27a total	and line 27b total			ightharpoons	27d	N/A	_
9	Public support (line 27c total minus line 27d total))			>	27e	N/A	_
:	Total support for section 509(a)(2) test Enter amount	ount on line 23, column (e)	▶ 27f	N/A				
,	Public support percentage (line 27e (num	erator) divided by line 27f (denominator))		>	27g	N/A %	,

Unusual Grants: For an organization described in line 10, 11, or 12 that received any unusual grants during 1999 through 2002, prepare a list for your records to show, for each year, the name of the contributor, the date and amount of the grant, and a brief description of the nature of the grant. Do not file this list with your return. Do not include these grants in line 15.

Investment income percentage (line 18, column (e) (numerator) divided by line 27f (denominator))

NONE 323121 12-05-03

27h

Schedule A (Form 990 or 990-EZ) 2003 HEALTH FOUNDATION, INC.

Part V Private School Questionnaire (See page 7 of the instructions.)

'(To be completed ONLY by schools that checked the box on line 6 in Part IV)

29	Does the organization have a racially nondiscriminatory policy toward students by statement in its charter, bylaws, other governing		Yes	No
	instrument, or in a resolution of its governing body?	29		
30	Does the organization include a statement of its racially nondiscriminatory policy toward students in all its brochures, catalogues,			
	and other written communications with the public dealing with student admissions, programs, and scholarships?	30		
31	Has the organization publicized its racially nondiscriminatory policy through newspaper or broadcast media during the period of	}		
	solicitation for students, or during the registration period if it has no solicitation program, in a way that makes the policy known			
	to all parts of the general community it serves?	31		
	If "Yes," please describe, if "No," please explain. (If you need more space, attach a separate statement.)	_		
		_		
32	Does the organization maintain the following:	_		ŀ
a	Records indicating the racial composition of the student body, faculty, and administrative staff?	328		
b	Records documenting that scholarships and other financial assistance are awarded on a racially nondiscriminatory basis?	32b		
C	Copies of all catalogues, brochures, announcements, and other written communications to the public dealing with student	222		1
	admissions, programs, and scholarships? Convey of all meteral used by the programment of an its helicit contributions?	32c 32d		
a	Copies of all material used by the organization or on its behalf to solicit contributions? If you answered "No" to any of the above, please explain. (If you need more space, attach a separate statement.)	32u		
33	Does the organization discriminate by race in any way with respect to:	_		
a	Students' rights or privileges?	33a		
Ь	Admissions policies?	33b		
C	Employment of faculty or administrative staff?	33c		
d	Scholarships or other financial assistance?	33d		
е	Educational policies?	33e		
f	Use of facilities?	33f		
g	Athletic programs?	33g		
h	Other extracurricular activities?	33h		<u> </u>
	If you answered "Yes" to any of the above, please explain. (If you need more space, attach a separate statement.)			
34 a	Does the organization receive any financial aid or assistance from a governmental agency?			
b	Has the organization's right to such aid ever been revoked or suspended?	34b		
	If you answered "Yes" to either 34a or b, please explain using an attached statement.			ĺ
35	Does the organization certify that it has complied with the applicable requirements of sections 4.01 through 4.05 of Rev. Proc. 75-50,			
	1975-2 C.B. 587, covering racial nondiscrimination? If "No," attach an explanation	35		

Page 5

Part VI-A | Lobbying Expenditures by Electing Public Charities (See page 9 of the instructions)

NT	1	λ	
v	•	А	

(To be completed ONLY by an eligible organization that filed Form 5768)

	_	(10 bo completed offer b)	an engible organization that nice roth of cor				
Che	ck ▶ a	If the organization belon	gs to an affiliated group. Check	b 🗀	ıf you che	cked "a" and "limited contr	ol" provisions apply.
			Lobbying Expenditures tures' means amounts paid or incurred.)			(a) Affiliated group totals	(b) To be completed for ALL electing organizations
						N/A	
36	Total lobb	iving expenditures to influence	public opinion (grassroots lobbying)		36	•	
37		. • .	a legislative body (direct lobbying)		37		
38		ying expenditures (add lines 3	• • • • • • • • • • • • • • • • • • • •		38		
39		mpt purpose expenditures	•		39		
40	Total exer	npt purpose expenditures (add	l lines 38 and 39)		40		
41	Lobbying	nontaxable amount. Enter the	amount from the following table -				
	if the amo	ount on line 40 is -	The lobbying nontaxable amount is -				
	Not over \$5	00,000	20% of the amount on line 40		۱ ا		
	Over \$500,0	000 but not over \$1,000,000	\$100,000 plus 15% of the excess over \$500,000				
	Over \$1,000	0,000 but not over \$1,500,000	\$175,000 plus 10% of the excess over \$1,000,000		41		
	Over \$1,500	0,000 but not over \$17,000,000	\$225,000 plus 5% of the excess over \$1,500,000				
	Over \$17,00	000,000	\$1,000,000		丿		
42	Grassroot	ts nontaxable amount (enter 25	5% of line 41)		42		
43	Subtract I	ine 42 from line 36. Enter -0- i	f line 42 is more than line 36		43		
44	Subtract I	line 41 from line 38. Enter -0- i	f line 41 is more than line 38		44	7-1-2782	
	Caution:	If there is an amount on eit	ther line 43 or line 44, you must file Form 4	720			

4-Year Averaging Period Under Section 501(h)

(Some organizations that made a section 501(h) election do not have to complete all of the five columns below. See the instructions for lines 45 through 50 on page 11 of the instructions.)

	Lobbying Expenditures During 4-Year Averaging Period						
Calendar year (or fiscal year beginning in)	(a) 2003	(b) 2002	(c) 2001	(d) 2000	(e) Total		
45 Lobbying nontaxable amount					(
46 Lobbying ceiling amount (150% of line 45(e))		٠	λ ,		(
47 Total lobbying expenditures					(
48 Grassroots nontaxable amount					(
49 Grassroots ceiling amount (150% of line 48(e))		*>	,		(
50 Grassroots lobbying expenditures	•						

Part VI-B Lobbying Activity by Nonelecting Public Charities

(For reporting only by organizations that did not complete Part VI-A) (See page 12 of the instructions.)

N/A

During the year, did the organization attempt to influence national, state or local legislation, including any attempt to influence public opinion on a legislative matter or referendum, through the use of:

- a Volunteers
- b Paid staff or management (Include compensation in expenses reported on lines c through h.)
- c Media advertisements
- d Mailings to members, legislators, or the public
- e Publications, or published or broadcast statements
- f Grants to other organizations for lobbying purposes
- g Direct contact with legislators, their staffs, government officials, or a legislative body
- h Rallies, demonstrations, seminars, conventions, speeches, lectures, or any other means
- $i \quad \text{Total lobbying expenditures (Add lines} \, c \, \, \text{through} \, \, h.)$
- If "Yes" to any of the above, also attach a statement giving a detailed description of the lobbying activities.

Yes	No	A	mount
\vdash		-	0.

323141 12-05-03

' Schedule A (Form 990 or 990-EZ) 2003 HEALTH FOUNDATION, INC.

Part \		garding Transfers To an zations (See page 12 of the inst		d Relationships With Noncharita	able		
51 Did		firectly or indirectly engage in any of		er organization described in section			
		section 501(c)(3) organizations) or		-			
		ganization to a noncharitable exemp		•		Yes	No
(i) Cash	•	-		51a(i)		Х
(ii) Other assets				a(ii)		X
b Ot	her transactions:						
(i) Sales or exchanges of asse	ets with a noncharitable exempt orga	inization		b(i)		X
(ii) Purchases of assets from a	a noncharitable exempt organization			b(ii)		X
(iii) Rental of facilities, equipme	ent, or other assets			b(iii)		X
(iv) Reimbursement arrangeme	ents			b(iv)	Х	
(v) Loans or loan guarantees				b(v)		X
•	•	r membership or fundraising solicita			b(vi)	X	-
		, mailing lists, other assets, or paid 6			С	X	<u> </u>
go	ods, other assets, or services	re is "Yes," complete the following so s given by the reporting organization ment, show in column (d) the value o	n. If the organization receive				
(a)	(b)	(c)	51 till goods, other decete, e	(d)			
Line no.	Amount involved	Name of noncharitable ex	kempt organization	Description of transfers, transactions, and st			nents
				REIMBURSEMENTS FOR O			
D T17		AMEDICAN KENNER	OI IID	POCKET EXPENDITURES	ON R	<u>EHA</u>	<u>.LF</u>
B-IV		AMERICAN KENNEL	СТОВ	OF FOUNDATION.			
				ADMINISTER PENSION A	M CIM	דמים	CA
B-VI		AMERICAN KENNEL	CLUB	PLAN FOR FOUNDATION.	ND M	<u> </u>	<u>CA</u>
D VI		AMBRICAN RUMNUD	СПОВ	THIN TOK TOOKBITTON:			
C		AMERICAN KENNEL	CLUB	SHARE MAILING LIST.			
				DONATED SPACE AND SE			
	42.075	AMEDICAN KENNIEL	OT IID	(PAYROLL, HUMAN RESON	<u>URCE</u>	4	
C	42,075.	AMERICAN KENNEL	СТОВ	ADMIN. SERVICES)			
		+			<u> </u>		
					••		
Co	the organization directly or in ode (other than section 501(c "Yes," complete the following	(3)) or in section 527?	one or more tax-exempt or	ganizations described in section 501(c) of the	Yes] No
	(a Name of or	1)	(b) Type of organization	(c) Description of relationshi	D		
		3	- 3,1	SHARE FACILITIES AND	<u> </u>		
			7	EQUIPMENT, PERSONNEL	SER	VIC	'판'S
AMER	ICAN KENNEL C	T.IIR	501(C)(4)	DONATED (SEE ABOVE)	DER	VIC	<u></u>
THILL	TOTAL KUMMUU (<u>,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,</u>	301(0)(1)	DOMITED (DEE 1150VE)			
	-						
					·		
							
		· · · · · · · · · · · · · · · · · · ·					
			-				
			ļ				

323151 12-05-03

FORM 990 GAIN (LOSS) FR	OM PUBLICLY T	RADED SECURIT	IES S	TATEMENT 1
DESCRIPTION	GROSS SALES PRICE	COST OR OTHER BASIS	EXPENSE OF SALE	NET GAIN OR (LOSS)
	- DALLED TRICE			——————————————————————————————————————
CD MATRIX 1.35% 9/4/03	75,000.	75,000.	0.	0.
CD STERLING 1.30%	73,000.	73,000.	0.	0.
8/26/03	85,000.	85,000.	0.	0.
EATON VANCE WW				
HEALTH & SCIENCE	3,611.	3,500.	0.	111.
FED. HOME LOAN BANK	105.000	400.040	•	4 040
5.575% 9/2/03	125,000.	129,848.	0.	<4,848.>
FED. HOME LOAN BANK 6.03% 3/27/09	35,000.	35,000.	0.	0.
FED. HOME LOAN BANK	33,000.	33,000.	0.	0.
6.57% 7/25/11	60,000.	60,019.	0.	<19.>
GREENPOINT FINANCIAL	00,000.	00,013.	•	(1)
CORP.	803.	833.	0.	<30.>
WRIGLEY WM JR CO	2,706.	2,620.	0.	86.
CATS 5/15/03	73,000.	73,000.	0.	0.
CD IBJ WHTHL 4.5%				
7/7/03	50,000.	50,000.	0.	0.
EXCELSIOR VAL &				
RESTRUCT	22,777.	30,746.	0.	<7,969.>
PBHG LARGE CAP VALUE	4,040.	5,271.	0.	<1,231.>
PBHG LARGE CAP VALUE	41,210. 12,429.	47,374.	0. 0.	<6,164.>
S&P 500 BARVAL INDEX WARNER LAM. 5.75%	12,429.	18,435.	0.	<0,000.2
1/15/03	50,000.	50,266.	0.	<266.>
CD SCHWAB VAR RATE	50,000.	50,000.	0.	0.
CD SCHWAB VAR RATE	35,000.	35,000.	0.	0.
FHLB 3.2% 9/12/06	150,000.	150,000.	0.	0.
FHLB 5.10% 3/13/07	95,000.	95,000.	0.	0.
FHLB 5.29% 3/26/07	40,000.	40,000.	0.	0.
FHLB 6.57% 7/25/11	40,000.	40,013.	0.	<13.
FREDDIE MAC	65,201.	75,720.	0.	<10,519.>
JOHN HANCOCK FINL			_	
SERVICES	18,731.	24,702.	0.	<5,971.>
MBNA CORP	25,926.	24,943.	0.	983.
NORTH FORK BANCORP	16,439.	21,025.	0.	<4,586.>
PBHG LARGE CAP VALUE	36,402.	42,103.	0.	<5,701.>
PHOENIX CAP W. MARKET NEUTRAL	71,282.	75,000.	0.	<3,718.>
SEARS 7.5% 1/15/13	23,767.	20,000.	0.	3,767.
SICOR INC.	24,259.	20,030.	0.	4,229.
TWEEDY BROWN GLOBAL	21,233.	20,030*	•	.,445
VALUE	33,422.	49,651.	0.	<16,229.
VERIZON	•	•		•
COMMUNICATIONS	30,106.	48,334.	0.	<18,228.
FHLB 3.2% 9/12/06	100,000.	100,000.	0.	0.
TO FORM 990, PART I, LINE 8	1,496,111.	1,578,433.	0.	<82,322.>

FORM 990 GAIN	(LOSS) FROI	M SALE OF OTH	HER ASSETS	STA	TEMENT 2
DESCRIPTION		DATE ACQUIF			
SALE OF FIXED ASSETS				PURCH	ASED
NAME OF BUYER	GROSS SALES PRICE	COST OR OTHER BASIS	EXPENSE OF SALE	DEPREC	NET GAIN OR (LOSS)
	3,334.	2,000.	0.	0.	1,334.
DESCRIPTION		DATE ACQUIF			
RETIREMENT OF FIXED ASS	SETS			PURCH	ASED
NAME OF BUYER	GROSS SALES PRICE	COST OR OTHER BASIS	EXPENSE OF SALE	DEPREC	NET GAIN OR (LOSS)
	0.	20,837.	0.	0.	<20,837.>
TO FM 990, PART I, LN 8	3,334.	22,837.	0.	0.	<19,503.>
FORM 990	SPECIAL E	VENTS AND ACT	PIVITIES	STA	TEMENT 3
DESCRIPTION OF EVENT	GROSS RECEIPT	CONTRIBUT	. GROSS REVENUE	DIRECT EXPENSES	NET INCOME
GALA 2003 - ORLANDO FLA (DINNER, AUCTION, RAFFI 2003 CANCER SYMPOSIUM,		1. 120,616	. 49,075.		49,075.
SEATLLE, WASHINGTON IKC BANQUET & RAFFLE SPECIAL EVENTS - OTHER	17,45 6,87				4,950. 6,875. 0.
TO FM 990, PART I, LINE	9 194,01	6. 133,116.	60,900.	-	60,900.

FORM 990 OTHER C	HANGES IN NET A	SSETS OR FUND	BALANCES	STATEMENT	4
DESCRIPTION				TNUOMA	
UNREALIZED APPRECIATION	ON PORTFOLIO		•	421,90	7.
TOTAL TO FORM 990, PART	I, LINE 20			421,90	7.
FORM 990	OTHER	REXPENSES		STATEMENT	5
	(A)	(B) PROGRAM	(C) MANAGEMENT	(D)	
DESCRIPTION	TOTAL	SERVICES	AND GENERAL	FUNDRAISING	G
PROFESSIONAL FEES MARKETING, ADVERTISING, ANNUAL	173,449.	45,943.	115,288.	12,21	8.
REPORT	43,605.	1,659.	21,974.		
MEMBERSHIP EXPENSES WEBSITE DESIGN AND	4,547.		2,171.	2,37	6.
EXPENSES	3,385.	2,629.	690.	6	6.
NEW DEVELOPMENT EXPENSES PROMOTIONAL ITEMS	139.		114.	2	5.
PURCHASED	8,083.			8,08	
MISCELLANEOUS RELOCATION EXPENSE, NET OF AKC \$70,000	33,057.	4,357.	25,147.	3,55	3.
REIMBURSEMENT	92,225.		92,225.		
TOTAL TO FM 990, LN 43	358,490.	54,588.	257,609.	46,29	 3.

	•						_
FORM 990	STATEMENT OF	ORGANIZATION'S	PRIMARY	EXEMPT	PURPOSE	STATEMENT	6
		PART I	II				

EXPLANATION

THE ORGANIZATIONAL EXEMPT PURPOSE IS TO SUPPORT BASIC AND APPLIED HEALTH PROGRAMS WITH EMPHASIS ON CANINE GENETICS TO IMPROVE THE QUALITY OF LIFE FOR DOGS AND THEIR OWNERS.

FORM 9'90	STATEMENT	OF PROGRAM	SERVICE	ACCOMPLISHMENTS	STATEMENT	7
DESCRIPTION (F PROGRAM SER	VICE ONE	 			

THE FOUNDATION FUNDS RESEARCH AND SUPPORTS CANINE HEALTH SCIENTISTS AND PROFESSIONALS IN THEIR EFFORTS TO STUDY THE CAUSES AND ORIGINS OF CANINE DISEASES AND AFFLICTIONS AND TO FORMULATE EFFECTIVE TREATMENTS. SEE ATTACHMENTS ENTITLED "GRANTS FOR RESEARCH ON SPECIFIC CANINE DISEASES AND FOR RESEARCH IN SPECIFIC BREEDS OF DOG".

			GRAI	NTS	EXPENSES	
TO FORM 990, PART III, LINE A			1,5	70,391.	1,981,91	19.
NON-GOVE	RNMENT SECU	RITIES		S	TATEMENT	8
CORPORATE STOCKS	CORPORATE BONDS	PUBL TRA	ICLY DED	OTHER SECURITIES	TOTAL NON-GOV' SECURITIE	
245,131.	245,969.	1,959	,960.		1,959,96 245,96 245,13	69.
245,131.	245,969.	1,959	,960.		2,451,06	50.
GOV	ERNMENT SEC	URITIE	S	s	TATEMENT	9
	U.S. GOVERNM	ENT			TOTAL GOV'	
ATIONS	677	,347.			677,34	47.
NE 54, COL E	677	,347.			677,34	47.
	NON-GOVE CORPORATE STOCKS 245,131. GOV	CORPORATE CORPORATE BONDS 245,969. 245,131. 245,131. 245,969. GOVERNMENT SEC U.S. GOVERNM GOVERNM 677	NON-GOVERNMENT SECURITIES CORPORATE CORPORATE TRA STOCKS BONDS SECUR 1,959 245,969. 245,131. 245,131. 245,969. GOVERNMENT SECURITIE U.S. GOVERNMENT GOVERNMENT 677,347.	NON-GOVERNMENT SECURITIES CORPORATE CORPORATE PUBLICLY TRADED SECURITIES 1,959,960. 245,131. 0. 245,131. 245,969. 1,959,960. GOVERNMENT SECURITIES U.S. STAGOVERNMENT LOCAL GATIONS 677,347.	NON-GOVERNMENT SECURITIES CORPORATE CORPORATE TRADED OTHER STOCKS BONDS SECURITIES SECURITIES 1,959,960. 245,131. 0. 245,131. 245,969. 1,959,960. GOVERNMENT SECURITIES U.S. STATE AND GOVERNMENT LOCAL GOV'T GATIONS 677,347.	NON-GOVERNMENT SECURITIES STATEMENT

FORM 990 OTHER INVESTMEN	ITS	STATEMENT	10
DESCRIPTION	VALUATION METHOD	AMOUNT	
CERTIFICATES OF DEPOSIT	COST	424,47	2.
TOTAL TO FORM 990, PART IV, LINE 56, COLUMN E	3	424,47	2.
FORM 990 OTHER ASSETS	3	STATEMENT	11
DESCRIPTION		AMOUNT	
CHARITABLE REMANDER ANNUITY TRUST RECEIVABLE DIVIDEND & INTEREST RECEIVABLE		78,49 10,21	
TOTAL TO FORM 990, PART IV, LINE 58, COLUMN E	3	88,71	1.
FORM 990 OTHER REVENUE NOT INCLUDED	O ON FORM 990	STATEMENT	12
DESCRIPTION		AMOUNT	
AKC RELOCATION EXPENSE REIMBURSEMENT		70,00	0.
TOTAL TO FORM 990, PART IV-A		70,00	0.
FORM 990 OTHER EXPENSES NOT INCLUDE	ED ON FORM 990	STATEMENT	13
DESCRIPTION		AMOUNT	
RELOCATION EXPENSE (REIMBURSED BY AKC) LOSS ON RETIREMENT OF FIXED ASSETS		70,00 20,83	
		90,83	

FORM 990 OTHER R	EVENUE	INCLUDED ON FOR	м 990	STAT	EMENT	14
DESCRIPTION				A	MOUNT	
LOSS ON RETIREMENT OF FIXED	ASSETS				<20,8	37.>
TOTAL TO FORM 990, PART IV-	A				<20,8	37.>
		OFFICERS, DIRE D KEY EMPLOYEES		STAT	EMENT	15
NAME AND ADDRESS		TITLE AND AVRG HRS/WK			EXPEN	
DR. SHELDON B. ADLER C/O AKC CANINE HEALTH FOUND INC. P.O. BOX 37941 RALEIGH, NC 27627	ATION,	0	0.	0.		0.
LEE ARNOLD C/O AKC CANINE HEALTH FOUND INC. P.O. BOX 37941 RALEIGH, NC 27627	ATION,	SECRETARY 0	0.	0.		0.
ALEXANDER F. DRAPER C/O AKC CANINE HEALTH FOUND INC. P.O. BOX 37941 RALEIGH, NC 27627	ATION,	TREASURER (THR	U 3/9/03)	0.		0.
ELYZABETH HIGGINS C/O AKC CANINE HEALTH FOUND INC. P.O. BOX 37941 RALEIGH, NC 27627	ATION,	EMERITUS DIREC	TOR	0.		0.
ROBERT L. KELLY C/O AKC CANINE HEALTH FOUND INC. P.O. BOX 37941 RALEIGH, NC 27627	ATION,	ACTING TREASUR	ER 3/20/03 0.	0.		0.
CATHERINE BELL C/O AKC CANINE HEALTH FOUND INC. P.O. BOX 37941	ATION,	SECOND VICE PR	ESIDENT	0.		0.

RALEIGH, NC 27627

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Ł	4	_	•	×	- 1	٠.	×	- 1	4
_	•		J	v	-	•	v	_	_

AMERICAN KENNEL CLUB CANINE HEAD	LTH FOUND		13-38	13813
C/O AKC CANINE HEALTH FOUNDATION,				
INC. P.O. BOX 37941 RALEIGH, NC 27627	57,.	115.	14,280.	0.
DR. ASA MAYS C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.
JOHN A. STUDEBAKER C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627		0.	0.	0.
	DIRECTOR EMERITUS			
C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	U	0.	0.	0.
HOWARD FALBERG C/O AKC CANINE HEALTH FOUNDATION,		3)		
INC. P.O. BOX 37941 RALEIGH, NC 27627	·	0.	0.	0.
DR. C. CRESTON FARROW C/O AKC CANINE HEALTH FOUNDATION,				
INC. P.O. BOX 37941 RALEIGH, NC 27627		0.	0.	0.
WAYNE FERGUSON C/O AKC CANINE HEALTH FOUNDATION,		/9/03		
INC. P.O. BOX 37941 RALEIGH, NC 27627	·	0.	0.	0.
MYRLE HALE C/O AKC CANINE HEALTH FOUNDATION,	0			
INC. P.O. BOX 37941 RALEIGH, NC 27627		0.	0.	0.
SUSAN LACROIX HAMIL C/O AKC CANINE HEALTH FOUNDATION,	0			
INC. P.O. BOX 37941 RALEIGH, NC 27627		0.	0.	0.
BRUCE ANDREW KORSON C/O AKC CANINE HEALTH FOUNDATION,	0			
INC. P.O. BOX 37941 RALEIGH, NC 27627		0.	0.	0.

AMERICAN KENNEL CLUB CANINE HEAI	TH FOUND		13-38	13813
PROF. IRIS C. LOVE C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.
DR. WILLIAM R. NEWMAN C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.
DR. ROBERT HRITZO C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	DIRECTOR EMERITUS 0	0.	0.	0.
PAMELA STEPHENS BUCKLES C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.
DR. DUANE BUTHERUS C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.
PAUL A. CAILLAUD C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.
THOMAS L. MILLNER C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.
DR. WILLIAM TRUESDALE C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.
MARY EDWARDS HAYES C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.
KAREN T. LEFRAK C/O AKC CANINE HEALTH FOUNDATION, INC. P.O. BOX 37941 RALEIGH, NC 27627	0	0.	0.	0.

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C/O AKC CANINE HEALTH FOUNDATION, 40 INC. P.O. BOX 37941 RALEIGH, NC 27627 STEVE REMSPECHER C/O AKC CANINE HEALTH FOUNDATION, 0	86,346.	25,322.	33,952.
- · - · - · - · - · · · · · · · · ·			
INC. P.O. BOX 37941 RALEIGH, NC 27627	0.	0.	0.
TOTALS INCLUDED ON FORM 990, PART V	143,461.	39,602.	33,952.

PART VI, LINE 90

STATES

NY, CA, CT, IL, MI, NJ, NC, PA, OH, MN, KS, MS, VA, SC, OK, AR, AL, WA, AZ, GA, MA, MO, NH, RI, WV, OR, WI, ME, UT, DC, NM, TN, ND, MD, AK, KY, FL, IN

SCHEDULE A

STATEMENT REGARDING ACTIVITIES WITH
SUBSTANTIAL CONTRIBUTORS, TRUSTEES, DIRECTORS,
CREATORS, KEY EMPLOYEES, ETC,.
PART III, LINE 2

STATEMENT 3

17

THE FOUNDATION REIMBURSES IT'S OFFICERS AND DIRECTORS FOR OUT OF POCKET EXPENDITURES MADE ON BEHALF OF THE FOUNDATION UPON RECEIPT OF AN ITEMIZATION OF SUCH EXPENDITURES.

AMERICAN KENNEL CLUB CANINE HEALTH FOUNDATION, INC. FEDERAL FORM 990, PAGE 3, LINES 57(A) & 57(B) STATEMENT:

NOTE 3 - FURNITURE, FIXTURES AND EQUIPMENT Furniture, fixtures and equipment consisted of the following

Furniture, fixtures and equipment, net	\$ 76,790	\$ 44,177	
Less accumulated depreciation	(15,381)	(90,277)	
Furniture, fixtures and equipment	\$ 92,171	\$134,454	
•	2003	2002	

American Kennel Club Canine Health Foundation, Inc. Attachment to Form 990, Year ended December 31, 2003 Confirmed Copy of Amended Bylaws of the American Kennel Club Canine Health Foundation, Inc. as of December 31, 2003.

To whom it may concern,

I confirm that the amended Bylaws attached hereto are complete and accurate copies of the original documents.

Bx: John A. Studebaker, President

Daked:

5/6/64



(A New York Not-for-Profit Corporation)

BYLAWS

ARTICLE I NAME

The name of the corporation is American Kennel Club Canine Health Foundation. The corporation shall hereinafter in these Bylaws be referred to as the "Corporation."

ARTICLE II OFFICE

The principal office of the Corporation shall be located in the City of New York, County of New York, State of New York, or such other location as the Board of Directors may designate.

ARTICLE III PURPOSES OF THE CORPORATION

The purpose of the Corporation is to further the knowledge of canine diseases and health care by clinical study, laboratory research, the sponsorship of educational programs, and taking any and all lawful steps in furtherance of this purpose, consistent with the Corporation's Certificate of Incorporation.

ARTICLE IV MEMBERS

The Board of Directors may establish categories of non-voting membership from individuals and organizations who indicated their interest in the purposes and programs of the foundation and who pay the appropriate dues. Rights, privileges and dues of such members may vary from category to category and shall be determined by the Board of Directors.

ARTICLE V BOARD OF DIRECTORS

Section 1. Qualification of Directors. Each director shall be at least eighteen (18) years of age.

Section 2. <u>Powers and Duties.</u> The Board of Directors shall have the general power and responsibility to control and manage the business, affairs and property of the Corporation, subject to applicable law and the Corporation's Certificate of Incorporation. It shall have full power, by majority vote of the directors present and voting at any duly constituted meeting, to adopt rules and regulations governing the action of the Board of Directors.

Specific notice regarding rules and regulations to be considered at a regular meeting of the Board of Directors need not be given.

The Board of Directors shall have full authority with respect to the distribution and payment of the monies received by the Corporation from time to time.

Section 3. Number, Election, Term of Office and Removal. The number of directors shall be not fewer than five (5), the number to be fixed from time to time by resolution of the Board adopted by the affirmative vote of a majority of the Board of Directors present and voting. The initial directors shall be the persons named in the Articles of Incorporation. The regular term of office shall be four years. Board members shall serve a maximum of three consecutive four year terms of office. One class shall be elected to regular terms at each annual meeting. All elections shall be by ballot. Mail ballots may be used if approved at least thirty days in advance by the affirmative vote of a majority of the entire Board of Directors. Directors whose term or class expire shall be elected at the annual meeting of the Board of Directors by the affirmative vote of a majority of the Directors present and voting, and each shall continue in office until the next annual meeting of the Board of Directors and until his or her successor shall have been elected and qualified or until his or her earlier death, resignation or removal.

Vacancies on the Board of Directors, for whatever reason, may be filled at any regular meeting of the Board, by a majority of the Board of Directors present and voting providing that prior notice has been given of such intent.

Any director may be removed at any time for cause providing prior notice has been given to the Board and to the Director a minimum of one month and a maximum of three months prior to such a vote. On December 9, 2001, the number of Directors was fixed at twenty-four.

Section 4. <u>Annual Meeting</u>; Notice. The annual meeting of the Board of Directors shall be held at the principal office of the Corporation or at such other place as the Board of Directors shall designate on such day generally in March, but not later than June and no more than six (6) months after the end of the Corporation's most recent fiscal year as the Board of Directors shall designate. Notice of the time, place and purposes of such annual meeting shall be given by the Secretary personally, by telephone or facsimile, or by mailing a copy thereof by first class mail or delivering the same to each director not less than thirty (30) days before such annual meeting.

Section 5. Other Meetings; Notice. Regular meetings shall be held in each calendar quarter and notice thereof shall be delivered to each director not less than seven (7) days before each such meeting. Other meetings of the Board of Directors may be called by the President or by any director upon verbal or written demand of not less than one-fourth of the entire Board of Directors, with such meeting to be held at the principal office of the Corporation or at such other place as may be designated in the notice of such meeting. Notice of the time, place and purposes of any such meeting shall be given by the Secretary personally, by telephone or facsimile, or by mailing a copy thereof by first class mail or delivering the same to each director not less than seven (7) days before such meeting.

- Section 6. <u>Waiver of Notice of Meeting</u>. Notice of any meeting of the Board of Directors may be waived orally or in writing, before or after the meeting. Attendance of any meeting without protest regarding defects in notice of any meeting or written approval of the minutes of any meeting shall be equivalent to waiver of notice thereof.
- Section 7. (a) <u>Action Without a Meeting</u>. Any action permitted to be taken by the Board of Directors may be taken without a meeting if all members of the Board of Directors consent verbally or in writing to the adoption of a resolution authorizing the action. The resolution and any written consents thereto by the members of the Board of Directors shall be filed with or recorded in the minutes or the proceedings or the Board of Directors.
- **(b)** Meetings by Conference Telephone. The members of the Board of Directors or any committee thereof may participate in a meeting of such board or committee by means of teleconferencing or similar communications equipment by means of which all persons participating in the meeting can communicate with each other and such participation shall constitute presence in person at such meeting.
- Section 8. Quorum; Adjustment of Meetings. At all meetings of the Board of Directors, a majority of the entire board, (but no less than two (2) members) of the entire board shall constitute a quorum for the transaction of business. In the absence of a quorum, a majority of the directors present may, without giving notice other than by announcement at the meeting, adjourn the meeting from time to time until a quorum is obtained. At any such adjourned meeting, at which a quorum is present, any business may be transacted which might have been transacted at the meeting as originally called. The act of a majority of the directors present at any meeting at which there is a quorum shall be the act of the Board of Directors unless a greater vote is required by law. Among the actions for which a greater vote is required by law are purchases, sales and mortgages of real property and leases of real property owned by the Corporation.
- Section 9. <u>Organization</u>. The President of the Corporation shall preside at all meetings of the Board of Directors or, in the absence of the President, the Vice-President, or in the absence of the President and the Vice-President, a Chairperson shall be chosen by a majority of the directors present. The Secretary of the Corporation shall act as Secretary at all meetings of the Board of Directors. In the absence of the Secretary, the person presiding at the meeting may appoint any person to act as Secretary of the meeting.
- **Section 10.** Compensation. No officer or director of the Corporation shall receive, directly or indirectly, any salary, compensation or emolument therefrom for his or her services as officer or director or in any other capacity except for expenses incurred at the request of the Executive Committee, unless authorized by the affirmative vote of a majority of the entire Board of Directors.
- **Section 11.** Resignation. Any director may resign at any time by giving written or oral notice to the President or the Corporation. Such resignation shall take effect at any time specified therein and, unless otherwise specified therein, the acceptance of such resignation shall not be necessary to make it effective. All Board members shall promptly be notified of any resignation.

- **Section 12.** <u>Director Emeritus</u>. The Board of Directors may designate a former member as Director Emeritus.
- **Section 13.** Parliamentary Authority. The rules contained in the current edition of Robert's Rules of Order Newly Revised shall govern the Corporation in all cases to which they are applicable and in which they are not inconsistent with this Constitution and Bylaws and any special rules or Order the Corporation may adopt.

ARTICLE VI EXECUTIVE COMMITTEE

- **Section 1. Qualifications.** The Executive Committee shall be composed of Officers and the immediate past president.
- **Section 2.** <u>Powers and Duties</u>. Between meetings of the Board of Directors, the Executive Committee shall have the authority to act on matters requiring attention but not in conflict with any action of the Board nor in any matter reserved by law to the Board of Directors.
- Section 3. <u>Meetings</u>. Meetings may be called by the President or at the request of any two members; two days' notice shall be given. A quorum shall be a majority.

The agenda shall be disseminated to all board members and any board member who wishes may be included in the meeting.

Section 4. Reporting. The committee shall report all actions within seven (7) days to the entire Board of Directors.

ARTICLE VII NOMINATING COMMITTEE

- Section 1. <u>Appointment and Duties</u>. Immediately following the annual election of officers, and prior to the next quarterly meeting, the President shall appoint an annual Nominating Committee of three or more Directors who are not officers to nominate candidates for vacancies on the Board created by the expiration of term and for positions as officers of the corporation to be filled at the annual meeting. Written nominations by the Committee shall be delivered to the Secretary 60 days prior to the Annual Meeting. The Secretary will forward to the Board of Directors the nominating committee's report. Additional written nominations signed by a minimum of two Board members for a position on the Board of Directors or for the position as officer, will close 21 days prior to the Annual meeting. There shall be no nominations made thereafter.
- Section 2. <u>Potential Candidates</u>. At any time, any two Directors may submit to the Nominating Committee a signed recommendation of a person they believe could be a new Director. The chair of the Committee shall determine the potential candidate's interest and willingness to serve and send to such person descriptive material regarding the Corporation and its work, a director's profile, and any other material it deems necessary, and shall ask for a brief biography of the candidate.

The Nominating Committee shall present its recommendations to the Board of Directors for its review, discussion and action.

Section 3. <u>Notification</u>. No person except the President shall notify a candidate of nomination or election to the Board of Directors.

ARTICLE VIII OFFICERS

Section 1. Officers. The officers of the Corporation shall be a President, a First Vice-President, a Second Vice-President, a Secretary, a Treasurer and such other officers, if any, as the Board of Directors may from time to time appoint or elect. One person may hold more than one office in the Corporation, except that one person may not hold both the offices of President and Secretary or President and Treasurer. No instrument required to be signed by more than one officer shall be signed by one person in more than one capacity. Except for the Executive Vice-President, only a director of the Corporation may serve as an officer.

Section 2. Election, Term of Office and Removal. The initial officers of the Corporation shall be selected by the initial directors of the Corporation. Thereafter, the officers of the Corporation shall be elected at the annual meeting of the Board of Directors immediately following the election of directors and shall hold office for two years. Officers may not serve more than two consecutive two-year terms in the same office. The maximum terms of office for the President, Second Vice President, and Secretary, will end in odd-numbered years, starting in 2007. The maximum terms of office for the First Vice President and Treasurer will end in even-numbered years, starting in 2008, resulting in an initial, one-time five-year term limit; the four-year term limit for these offices will apply thereafter. Any officer of the Corporation may be removed at any time, with or without cause, by the affirmative vote of a majority of the directors then in office.

Section 3. Other Agents. The Board of Directors may from time to time appoint such agents as it shall deem necessary, each of whom shall hold office at the pleasure of the Board of Directors, and shall have such authority, compensation, if any, as the Board of Directors may from time to time determine.

Section 4. <u>Vacancies</u>. Any vacancy in any office may be filled by the Board of Directors at any meeting. Any officer so elected shall hold office until the next annual meeting.

Section 5. <u>President; Powers and Duties</u>. The President shall be the chief executive officer of the Corporation and shall preside at all meetings of the Board of Directors. The President shall generally manage and supervise the affairs of the Corporation. The President shall keep the Board of Directors fully informed, and shall freely consult with them concerning the activities of the Corporation. The President shall present at the annual meeting of the Board of Directors a report complying with Section 519 of the New York Not-for-Profit Corporation Law, which report shall be filed with the records of the Corporation.

The President shall have the power to sign alone, unless the Board of Directors shall specifically require an additional signature, in the name of the Corporation all contracts authorized either generally or specifically by the Board of Directors. The President shall perform all duties incident to the office of President, subject to the direction of the Board of Directors and such other duties as shall from time to time be assigned to him or her by the Board of Directors.

Section 6. Secretary; Powers and Duties. The Secretary shall act as secretary of all meetings of the Board of Directors and shall keep the minutes of all such meetings in the books proper for that purpose and shall distribute copies of such minutes to all Directors within three weeks after each meeting. As a procedural, not as part of the bylaws, a draft is to be sent within one week of the meeting, by fax, with corrections/additions returned to the secretary again within another week by fax. The Secretary shall attend to the giving and serving of all notices of the Corporation. The Secretary shall have custody of the seal of the Corporation and shall affix the same to all instruments requiring it when authorized by the Board of Directors or President, and attest the same. The Secretary shall perform all duties incident to the office of the Secretary subject to the direction of the President and such other duties as shall from time to time be assigned to him or her by the President or by the Board of Directors.

Section 7. <u>First Vice-President</u>; <u>Powers and Duties</u>. The First Vice-President shall assume the powers and duties of the President in the absence, incapacity or death of the President.

Section 8. <u>Second Vice-President; Powers and Duties.</u> The Second Vice-President shall assume the powers and duties of the President in the absence, incapacity or death of the President and the absence, incapacity or death of the First Vice-President.

Section 9. Treasurer; Powers and Duties. The Treasurer shall have the custody of all funds, securities, evidences of indebtedness and other valuable documents of the Corporation, which may come into his or her hands. The Treasurer shall keep or cause to be kept complete and accurate accounts of receipts and disbursements of the Corporation, and shall deposit all moneys and other valuable effects of the Corporation in the name and to the credit of the Corporation in such banks or depositories as the Board of Directors may designate. Whenever required by the Board of Directors, the Treasurer shall render a statement of his or her accounts. The Treasurer shall at all reasonable times exhibit his or her books and accounts to any officer or director of the Corporation, and shall perform all duties incident to the office of Treasurer subject to the direction of the President and such other duties as shall from time to time be assigned to him or her by the President or by the Board of Directors. The Treasurer shall, if so required by the Board of Directors, give such security for the faithful performance of his or her duties as the Board of Directors may require.

ARTICLE IX ADVISORS

Section 1. Advisors. The Board of Directors may appoint from time to time any number of persons as advisors of the Corporation to act either singly or as a committee or committees. Each such advisor shall hold office at the pleasure of the Board of Directors and shall have only such authority or obligation as the Board of Directors may from time to time determine.

Section 2. <u>Compensation</u>. No advisor of the Corporation shall receive, directly or indirectly, any salary, compensation or emolument therefrom for any service rendered to the Corporation by such advisor, unless authorized by the concurring vote of two-thirds of all the directors then in office. No director or officer of the Corporation shall be eligible for appointment as a paid advisor.

ARTICLE X CONTRACTS, CHECKS, BANK ACCOUNTS, INVESTMENTS, ETC.

Section 1. Checks, Notes, Contracts, Etc. The Board of Directors is authorized to select such banks or depositories, as it shall deem proper for the funds of the Corporation. The Board of Directors shall determine who, if anyone, in addition to the President, the Secretary and the Treasurer, shall be authorized from time to time on the Corporation's behalf to sign checks, drafts or other orders for the payment of money, acceptances, notes or other evidences of indebtedness, to enter into contracts, or to execute and deliver other documents and instruments.

Section 2. <u>Investments.</u> The funds of the Corporation may be retained in whole or in part in cash or be invested and reinvested from time to time in such property, real, personal or otherwise, or stocks, bonds or other securities, as the Board of Directors in its discretion may deem desirable.

Section 3. Gifts. The Board of Directors may accept on behalf of the Corporation any contribution, gift, bequest, or devise for the general purposes, or for any special purpose, of the Corporation.

ARTICLE XI BOOKS

Section 1. <u>Books</u>. There shall be kept at the principal office of the Corporation correct books of account of the activities and transactions of the Corporation, including a minute book, which shall contain a copy of the Certificate of Incorporation, a copy of these Bylaws and all minutes of meetings of the Board of Directors.

ARTICLE XII CORPORATE SEAL

The seal of the Corporation shall be circular in form and shall bear the name of the Corporation and words and figures showing that it was incorporated in the State of New York in 1994.

ARTICLE XIII FISCAL YEAR

The fiscal year of the Corporation shall end with the thirty-first day of December of each year.

ARTICLE XIV INDEMNIFICATION

Section 1. <u>Indemnification</u>. The Corporation shall, to the fullest extent now or hereafter permitted by law, indemnify any person made, or threatened to be made, a party to any action, suit or proceeding by reason of the fact that he or she (or a person of whom he or she is the legal or personal representative or heir or legatee) is or was a director, officer, employee or other agent of the Corporation, or of any other organization served by him or her in any capacity at the request of the Corporation, against judgments, fines, amounts paid in settlement and reasonable expenses, including attorney's fees. Such right or indemnification shall be a contract right which may be enforced in any manner such person may elect.

Section 2. Other Indemnification Rights. Such right of indemnification shall not be exclusive of any other rights which those indemnified may have or hereafter acquire under any bylaws, agreements, resolution of directors, provisions of law or otherwise.

Section 3. <u>Insurance</u>. The Board of Directors shall have the power to authorize the Corporation to purchase and maintain insurance (i) to indemnify the Corporation against liability incurred by the Corporation in connection with the activities of the Corporation, (ii) to indemnify the Corporation for any express obligation which it incurs as a result of the indemnification of any person under the provisions of this Article, and (iii) to indemnify any person who is or was a director, officer or employee of the Corporation, or the legal representative for such a person, against all expenses, liability and loss incurred by or asserted against such person in such capacity of arising out of such status, whether or not the Corporation would have the power to indemnify such person.

Section 4. Amendments. The Board of Directors may from time to time adopt further bylaws with respect to indemnification permitted by the laws of the State of New York.

ARTICLE XV AMENDMENTS

These Bylaws or any part thereof may be amended or repealed at any meeting of the Board of Directors by the affirmative vote of a majority of the Board of Directors present and voting, provided that notice of intention to amend the Bylaws and the proposed changes shall have been contained in the notice of the meeting.



AMERICAN KENNEL CLUB

CANINE HEALTH FOUNDATION

Grants for research on Specific Canine Diseases and Research to Benefit Specific Breeds of Dogs

Grants for research in specific diseases are based on scientific merit and established significance of the disease in dogs. Grants for research to benefit specific breeds are also based on scientific merit and significance in a particular breed as identified by that breed's national Parent Club. Research for specific breeds falls under the Club Partnership Program, which outlines the availability of matching funds and the opportunities for clubs to open restricted funds with the AKC Canine Health Foundation.

Allergies

Active Grant No 2234 Basophil/Mast Cell Response to Lectins as a Predictor for Risk of Allergic Disease in Genetically Susceptible Dogs

Principal Investigator: Bruce Hammerberg, DVM, PhD; North Carolina State University Sponsor(s). American Belgian Tervuren Club, Inc., American Miniature Schnauzer Club, Inc., American Sealyham Terrier Club, Bedlington Terrier Club of America, Bichon Frise Club of America, Inc., Bull Terrier Welfare Foundation, Chow Chow Club, Inc., Dalmatian Club of America Foundation, Inc., French Bulldog Club of America, Irish Water Spaniel Club of America, Otterhound Club of America, Rhodesian Ridgeback Club of the United States, Scottish Terrier Club of America Health Trust Fund, Welsh Terrier Club of America, Inc., Westie Foundation of America, Inc.

Abstract: Atopic dermatitis or skin allergies is a chronic debilitating disease that is widely distributed among the breeds of dogs. This inherited disease is listed as a high research priority for the following breeds: Bichon Frise, Boston Terrier, Bull Terrier, Cairn Terrier, Dalmatian, Vizsla, Welsh Terrier and West Highland White Terrier. The skin mast cell and circulating basophil are the cells mainly responsible for itching and skin damage seen in atopic dermatitis. This laboratory has just recently discovered that mast cells from atopic dogs release significantly more of the inflammatory mediator, tumor necrosis factor alpha (TNF-\(\alpha\)), than normal dog mast cells when stimulated with lectins that bind glycoproteins on the surface of mast cells. If there is an inherited difference in how surface glycoproteins signal release of TNF-\(\alpha\), then knowledge of the molecular basis for this difference will lead to being able to identify dogs that will have a

higher risk of developing atopic dermatitis. To accomplish this, atopic and nonatopic dogs will be compared with regard to the identity and quantity of the cell surface glycoproteins on basophils that are responsible for signaling immediate TNF- α release stimulated by lectins.

Alternative and Complementary Medicine

Active Grant No. 2628. The Effects of Acupuncture, Electroacupuncture and Transcutaneous Cranial Electrical Stimulation on Isoflurane Requirements in Dogs: A Comparative Study Principal Investigator: Roman T. Skarda, DVM, PhD; Ohio State University Abstract: This randomized study should demonstrate that acupuncture, electroacupuncture (EA). and transcutaneous cranial electrical stimulation (TCES) significantly reduce the minimum alveolar concentration of isoflurane (MAC-ISO) and release the neurochemical beta-endorphin. to produce a sparing of isoflurane and post-anesthetic pain relief. Isoflurane is the most popular and commonly used inhalation anesthetic in dogs. Non-pharmacological therapy such as acupuncture, EA and TCES on MAC-ISO reduction and post-anesthetic analgesia in dogs, surprisingly, has not been investigated. A direct comparison of acupuncture, EA and TCES on isoflurane-sparing effects will demonstrate which of the three techniques is the most effective for reducing MAC-ISO. Electroacupuncture and TCES are expected to be more potent to reduce MAC-ISO than acupuncture. There are substantial benefits from minimizing the necessary concentration of isoflurane for general anesthesia, such as increased safety of anesthesia, reduced amount of waste anesthetic gases, minimal pollution of the environment, decreased toxic byproducts of anesthetic gases (carbon monoxide, inorganic fluoride), and reduced cost of inhalation anesthesia. We will determine the plasma concentration of beta-endorphins and respiratory and cardiovascular effects of acupuncture, EA and TCES in healthy dogs during and after administration of isoflurane.

Autoimmune Disease

Active Grant No. 2226: Characterizing the Inheritance of Addison's Disease and Linked DNA Markers

Principal Investigator Anita M. Oberbauer, PhD; University of California, Davis Sponsor(s): Anonymous, BEACON for Health, Bearded Collie Club of America, Poodle Club of America Foundation, Portuguese Water Dog Club of America, Inc., Portuguese Water Dog Foundation, Versatility in Poodles, Inc., Westie Foundation of America, Inc. Abstract: Addison's disease is a late onset disorder caused by the deterioration of the adrenal gland. Addison's occurs in the domestic dog at approximately 0.1 percent, with some breeds showing a greater prevalence. Notably, the Bearded Collie, the West Highland White Terrier, the Standard Poodle, the Portuguese Water Dog, and the Leonberger are considered to have unacceptable rates of Addison's disease. Breeders have noted a familial tendency of Addison's disease suggesting a genetic basis to the disorder. Our laboratory has determined that Addison's is highly heritable in Bearded Collies. Further, although Addison's is not fully governed by a single locus in the Bearded Collie, it does appear to be regulated by a single gene of large effect. The specific objectives of this study are to develop a genetic marker associated with an

Addison's locus in the Bearded Collie; such a genetic marker will provide a useful tool to aid breeders in making health-based breeding decisions. The second objective is to determine if Addison's disease in the Standard Poodle, West Highland White Terrier, Portuguese Water Dog and Leonberger also has a genetic basis and if so, whether there is a common genetic defect across all these breeds.

Pending Grant No. 225. Establishing a Genetic Linkage Between Addison's Disease and DNA Markers

Principal Investigator. Anita M. Oberbauer, PhD; University of California, Davis Abstract: Addison's disease is a late onset disorder caused by deterioration of the adrenal gland cortex. Although Addison's disease occurs in the general canine population, some breeds show a greater prevalence as noted by owners and breeders: Bearded Collies, Standard Poodles. Leonbergers, Portuguese Water Dogs, and West Highland White Terriers. We have demonstrated that for the above breeds for which we have sufficient data to analyze. Addison's disease is highly heritable. Statistical evaluation of the dogs' pedigrees suggests a single locus of large effect significantly influences the expression of Addison's in the Standard Poodle and that this locus acts as an autosomal recessive. Similar findings characterize Addison's for the Bearded Collie although the level of significance is less robust. Thus, the specific objectives of this grant are to expand our pedigree, phenotypic, and DNA databases for all possible Bearded Collies, Standard Poodles, Leonbergers, Portuguese Water Dogs and West Highland White Terriers as related to Addison's disease and to continue our genome scan of the DNA to identify a genetic marker linked to the single locus suggested by the pedigree analyses. We have made an effort to include dogs affected with Addison's and those that are unaffected to give us the greatest depth of pedigree information that will speed the genome search.

Active Grant No 2270 Mechanisms for Hypercoagulability in Immune Mediated Hemolytic Anemia and Early Disseminated Intravascular Coagulation Principal Investigator. Cynthia Otto, DVM, PhD; University of Pennsylvania Sponsor(s) Collie Health Foundation, Flat-Coated Retriever Foundation, Versatility in Poodles, Inc.

Abstract: The formation of excessive blood clots (thrombosis) is a major cause of mortality in a variety of diseases that affect purebred and mixed breed dogs. Two very common conditions are immune-mediated hemolytic anemia (IMHA; a disease in which the dog's immune system destroys its own red blood cells) and disseminated intravascular coagulation (DIC; a syndrome in which inflammation initiates systemic coagulation, with subsequent depletion of clotting factors and inhibitors and the ultimate development of bleeding). Once thromboses form, specialized or invasive procedures must be used to confirm their presence and define the extent of organ involvement. Even with definitive diagnosis, treatment is limited. This study is designed to test the ability of a specialized clotting test, thromboelastography, to identify increased coagulation (hypercoagulability) in dogs with IMHA, and early DIC associated with a primary diagnosis of pancreatitis. Through our studies of the mechanisms of hypercoagulability in IMHA and DIC, we can design specific treatment strategies for the prevention of the devastating syndromes of thromboembolic (inappropriate formation and migration of clots) disease in IHMA and DIC. The ability to diagnose animals at risk for the complications of hypercoagulable states is revolutionary and will change critical care practice and treatment of these potentially fatal syndromes.

Active Grant No. 2273: Pre-clinical Detection of Hypoadrenocorticism (Addison's Disease) in dogs: Development and Evaluation of Laboratory Techniques for the Diagnosis of Immune-Mediated Canine Adrenal Disease

Principal Investigator Markus Rick, VetMed; Michigan State University Sponsor(s) Anonymous, Bearded Collie Club of America, Portuguese Water Dog Club of America, Inc., Westie Foundation of America, Inc.

Abstract: The adrenals are hormone-producing glands located near each kidney. Their products are of such importance that, without them, an individual cannot survive. If the adrenals are damaged and not able to make sufficient hormones, a potentially life-threatening disease called hypoadrenocorticism arises. This disease is known as Addison's disease. Both humans and dogs can have this disease. It is most often caused by an immune reaction in which the body's own defense mechanisms destroy the adrenals. This process releases telltale antibodies into the circulation before severe damage occurs. We hope to be able to measure these antibodies in serum. With this test, we will be able to make an early diagnosis long before affected animals become ill. Such a test will also help us identify which breeds, families and ages are most at risk for this disease. With this information we can identify which individual animals require close monitoring for the development of the disease and what breeding strategies might help eradicate it. An average prevalence of 0.3 percent has been reported for dogs, however numerous breeds including Bearded Collie, West Highland White Terrier, Standard Poodle, Portuguese Water Dog, Leonberger, Great Dane, Airedale Terrier, Basset Hound, Wheaten Terrier and Rottweiler are reported to be at higher risk.

Behavior Disorders

Active Grant No. 1871: Bull Terrier Families Affected With Compulsive Tail Chasing Behavior: Behavioral Diagnosis, Pedigree Collection and DNA Isolation for Future Genetic and DNA Isolation for Future Genetic Mapping Studies

Principal Investigator: Alice Moon-Fanelli, PhD; Tufts University

Sponsor(s). Bull Terrier Welfare Foundation

Abstract: Spinning and tail chasing is the most common form of compulsive behavior shown by Bull Terriers. In a previous study, we defined the clinical signs, development, interaction with environmental, physiologic and psychological triggers, and response to pharmacological treatment in a sample population. Our pilot pedigree data for a large Bull Terrier family affected with tail chasing indicate that the disorder has a heritable component in this breed. Over the next two years, we will locate families of Bull Terriers affected with compulsive behavior. From this population, we will systematically collect detailed descriptive, medical historical and pedigree data for them and for two to three generations of relevant family members. We will collect blood samples from these affected Bull Terriers and their relatives, isolate the DNA and store the samples for genetic mapping studies to follow this project. The goal of these two projects is to identify chromosomal regions that contain susceptibility genes for compulsive behavior in Bull Terriers. Successful completion of both of these projects (DNA isolation and gene mapping) should have a significant impact on understanding the genetic basis of compulsive behavior in Bull Terriers. Our long-term goal is to use this information to contribute to the development of diagnostics, a treatment and a cure for the disease.

Active Grant No. 22111. Assessing Canine Temperament - Establishing and Validating a Physiological Measurement to Quantify Emotional Reactivity in the Canine Principal Investigator: Paul Mundell; Canine Companions for Independence Abstract: Each year behavioral problems stemming from fear, nervousness or aggression result in dogs being surrendered to animal shelters or euthanized. Unfortunately, these problematic behaviors are often not recognized until the dog is adopted into a home, bred, or placed in a working role. For assistance dog organizations, temperament problems account for nearly 70 percent of the dog failure rate. Our project begins with the finding that an important component of temperament is rooted in emotional reactivity. Our multi-stage project seeks to define both an objectively scored behavioral test and a physiological parameter, the dog's fecal cortisol profile (FCP) that can be used to describe and quantify canine emotional reactivity. Unlike other physiological indicators of stress, assaying the FCP is non-invasive and reflects adrenal activity over time. The FCP should therefore be less sensitive to episodic and circadian fluctuations in adrenal secretions and allow us a clearer picture of whether an animal is chronically hypersecreting due to a higher degree of emotional reactivity. The project makes use of the large canine population at Canine Companions for Independence, which ranges in age from newborns to seniors and includes purebred Labrador and Golden Retrievers.

Active Grant No. 2433. Final Testing, Completion and Publication of the Behavioral Assessment and Research Questionnaire (PennBARQ)

Principal Investigator: James Serpell, PhD; University of Pennsylvania

Sponsor(s): Westie Foundation of America, Inc.

Abstract: Canine behavior problems are a significant public health concern in the USA, as well as being a major threat to the welfare and survival of pet dogs. Despite the extent of the problems posed by canine behavior problems, relatively little is known about their overall prevalence in the pet dog population, or the reasons why they develop in some dogs but not in others. Efforts to understand the distribution and development of behavior problems are currently hampered by the fact that (a) canine behavior is difficult to observe and record in the home environment using conventional methods, and (b) there is no widely accepted "gold standard" system for describing and classifying these problems. The proposed study is seeking to overcome these difficulties by completing the development and validation of a new behavioral measurement instrument, the University of Pennsylvania Behavioral Assessment and Research Questionnaire (PennBARQ). This instrument shows considerable promise both as a research tool for studying canine behavioral problems at the population level, and as a reliable method for screening individual dogs for the presence of behavior problems.

Brain Disease

Active Grant No 2501: Transmission Analysis of Breed Specific Necrotizing Encephalitis in the Pug Dog: Pedigree Collection Phase

Principal Investigator Kimberly Greer, PhD; Texas A&M University

Sponsor(s) Pug Dog Club of America, Inc.

Abstract: The long-term goal of the proposed research is to understand the molecular genetics of breed specific necrotizing meningoencephalitis in the Pug (Pug Dog Encephalitis, or PDE), the etiology of which is currently unknown. However, the breed predilection of PDE strongly

suggests a genetic component. A necessary resource for this goal is detailed information regarding the incidence and inheritance of PDE. To this end, we propose collection of pedigree information and biological samples. We will identify informative pedigrees and collect blood/tissue samples, together with extensive laboratory and clinical information, from all available pedigree members. The clinical data are necessary for definitive diagnosis of PDE and analyses of pedigrees will allow determination of the mode of transmission. Once this is known, we will carry out whole genome scans to reveal genetic markers associated with PDE. Identification of markers linked to PDE will 1) allow development of marker-based tests for carrier detection and 2) facilitate positional cloning of the candidate gene, or genes, causative for the disease.

Active Grant No. 2631. Polymerase Chain Reaction for Herpesviruses From Paraffinized Brains and Fresh Tissues in Cases of Pug Dog Encephalitis

Principal Investigator: Scott J. Schatzberg, DVM, PhD; Cornell University

Sponsor(s): Pug Dog Club of America

Abstract: Pug Dog encephalitis (PDE) is a fatal brain disease of young Pugs for which the cause is currently unknown. Anecdotally, the disease is over-represented in certain families of Pugs, and most investigators believe that the Pug Dog has a genetic predisposition for the development of this condition. Interestingly, similar brain diseases have been described in a variety of other small breed dogs, most notably the Yorkshire Terrier and Maltese. As no definitive inheritance pattern has been identified in any breed, we believe that the cause is multifactorial. We hypothesize that PDE is secondary to an inherited genetic defect that causes an exaggerated, fatal immune response following a herpesvirus infection. We are most suspicious of a herpesvirus because the clinical signs and brain lesions seen in PDE are remarkably similar to those appreciated in people with herpes simplex virus encephalitis. We plan to use the polymerase chain reaction (PCR), a molecular biology tool, to study archival Pug brains (collected over the years from dogs that have died from PDE) for the presence of herpesvirus DNA. The identification of a herpesviral cause for PDE would revolutionize the way we diagnose and treat this fatal disease.

Active Grant No 2650: Transmission Analysis of Breed Specific Necrotizing Encephalitis in the Pug Dog

Principal Investigator: Keith E. Murphy, PhD; Texas A&M University

Sponsor(s): Pug Dog Club of America, Inc.

Abstract: The long-term goal of the proposed research is to understand the molecular genetics of breed specific necrotizing meningoencephalitis in the Pug (Pug Dog Encephalitis, or PDE), the etiology of which is currently unknown. However, the breed predilection of PDE strongly suggests a genetic component. A necessary resource for this goal is detailed information regarding the incidence and inheritance of PDE. To this end, we propose collection of pedigree information and biological samples. We will identify informative pedigrees and collect blood/tissue samples, together with extensive laboratory and clinical information, from all available pedigree members. The clinical data are necessary for definitive diagnosis of PDE and analyses of pedigrees will allow determination of the mode of transmission. Once this is known, we will carry out whole genome scans to reveal genetic markers associated with PDE. Identification of markers linked to PDE will 1) allow development of marker-based tests for

carrier detection and 2) facilitate positional cloning of the candidate gene, or genes, causative for the disease.

Cancer

Active Grant No. 2035. Transferrin Receptor Expression by Canine Brain Tumors Principal Investigator. Natasha Olby, PhD; North Carolina State University Sponsor(s) Golden Retriever Foundation

Abstract: Dogs that are not treated following diagnosis of a brain tumor survive an average of only two weeks. Treatment with either surgery and/or radiation can extend their survival to about ten months. Clearly there is a need for new approaches to treating brain tumors in dogs if survival is to be improved. In humans, malignant brain tumors are being treated successfully with toxins targeted specifically to a marker expressed by tumor cells: the transferrin receptor. In preliminary work, we have shown that untreated canine brain tumors also express this marker. The primary aim of this project is to establish whether brain tumors that have recurred following treatment with radiation or surgery still express the transferrin receptor. This information will allow us to determine whether transferrin receptor targeted toxins should be used as the primary form of treatment of brain tumors, or can be used to treat recurrent tumors following more conventional therapy. We hypothesize that brain tumors will express transferrin receptors at high levels after treatment and therefore that transferrin-linked therapy can be instituted in combination with more conventional therapy.

Active Grant No 2038T: The Molecular Cytogenetics of Canine Lymphosarcoma: Correlating Chromosomal Changes with Clinical Disease

Principal Investigator: Matthew Breen, PhD; North Carolina State University Sponsor(s): American Bullmastiff Association, Bull Terrier Welfare Foundation, Chinese Shar-Pei Club of America Charitable Trust, Flat-Coated Retriever Foundation, Newfoundland Club of America Charitable Trust, Pembroke Welsh Corgi Club of America, Rhodesian Ridgeback Club of the United States, Rottweiler Health Foundation

Abstract: Cancer kills. Twenty years ago, the diagnosis of lymphosarcoma (a tumor of the lymph glands) in humans was almost invariably fatal. However, with the development of improved means to sub-classify this neoplasm and the tailoring of therapies that are subtype-specific, more and more forms of lymphosarcoma are treatable. One of the most important means of sub-classification of human tumors is based on the identification of chromosome abnormalities. In the dog, lymphosarcoma comprises one in five malignancies; however, the extent and identity of chromosome aberrations is still unknown. This is largely because the chromosomes of dogs were extremely difficult to identify with confidence. Recently, we have developed a set of canine chromosome-specific reagents that allow us to identify conclusively every dog chromosome. We propose to use these reagents to identify the chromosome aberrations associated with dog lymphosarcoma and to investigate the correlation between these aberrations and the clinical disease. Such an approach offers a means to potentially sub-divide this diverse disease in dogs, thereby offering new information of diagnosis, prognosis and therapy. Identification of specific chromosome aberrations will also help to investigate the correlation between the genetic etiologies in dogs with those in humans.

Active Grant No. 2214T. Identification of a 5-10Mb BAC Set as a Cytogenetic Resource and for the Development of an Ordered CGH Microarray for Cancer Studies in the Dog Principal Investigator: Matthew Breen, PhD; North Carolina State University Sponsor(s): American Boxer Charitable Foundation, Nestle Purina PetCare Company Abstract: The study of aberrant chromosome structure has significantly increased our understanding of the cause and progression of human cancers. Many cancers are common to both dogs and humans, in part reflecting the high degree of similarity in their genetic material and in their environmental exposure to carcinogens. The extent and identity of chromosome aberrations associated with canine cancers, however, remains largely unknown. This is primarily due to the difficulty in identifying dog chromosomes by conventional means alone. Comparative genomic hybridization (CGH) is a technique that allows a comprehensive analysis of chromosome aberrations present within tumors. We have developed this technique for application to dog cancers and are obtaining valuable information. However, chromosome-based CGH is labor intensive, has a limited resolution and requires detailed knowledge of dog chromosomes. In this study we aim to identify set of large insert clones that are evenly spaced, at small intervals, along all dog chromosomes. These clones will be a very valuable resource for chromosome studies of dog cancers. The clones will be used to generate an ordered microarray to replace chromosome-based analyses. This will significantly and rapidly advance the study of canine cancer, leading to improved diagnosis and prognosis and thus health and welfare.

Active Grant No. 22101: Development of Anti-Canine II-2Ra Antibodies Using CpG Oligodeoxynucleotide Vaccination

Principal Investigator: Stuart Helfand, DVM; University of Wisconsin, Madison Sponsor(s): American Boxer Charitable Foundation, Golden Retriever Foundation, Nestle Purina PetCare Company

Abstract: Despite progress in treating canine lymphoma, most affected dogs eventually develop resistance to chemotherapy and succumb to their disease. In human medicine, the á subunit of the interleukin-2 receptor (IL-2Rά) has been developed as a target for Immunotherapy of chemoresistant lymphoma patients. Intensively studied as a molecule present on normal lymphocytes that are activated, IL-2Rá has recently been developed as a target on malignant lymphocytes that have been found to express this protein inappropriately. This has been accomplished using anti-IL-2Rά monoclonal antibody (Mab) to deliver cellular toxins or radioisotopes directly to the cancer cells, a strategy that overcomes drug resistance. We would now like to develop such an antibody against the canine Il-2Rá subunit that could benefit dogs with lymphoma. Our rationale for pursuing this approach stems from molecular data we generated indicating that a high percentage of canine lymphomas from Golden Retrievers and Rottweilers express Il-2Rá. Having synthesized canine recombinant (cr) IL-2Rá using the gene sequence we cloned, we are now ready to produce and validate Mabs against it. To accomplish these goals, 1) we will employ a novel immunization strategy to prepare murine anti-canine IL-2Rá Mabs, 2) develop a test using anti-canine IL-2Rά Mabs for the detection of IL-2Rά shed into the blood of Golden Retriever and Rottweiler lymphoma patients, and 3) screen lymphoma biopsies from Golden Retrievers and Rottweilers for IL-2Rά. Mabs generated to Canine IL-2Rά will be a powerful tool in the diagnosis, prognosis, and treatment of canine lymphoma.

Active Grant No. 2222 Molecular Control of Prostaglandin Synthesis in Canine Mammary Tumors

Principal Investigator: Monique Doré, DVM, PhD; University of Montreal Sponsor(s) Mid-Kentucky Kennel Club, Nestle Purina PetCare Company, Skye Terrier Foundation

Abstract: Tumors originating from the mammary gland represent the most frequent form of cancer in female dogs. However, very little information is known of the mechanisms involved in the development of this cancer. In dogs, the growth of mammary gland tumors is dependent on the age of the animal and on its hormones. Malignant mammary tumors can recur following surgical excision or can send cancerous cells to distant organs. Cyclooxygenase-2, an enzyme that is essential in the production of prostaglandins, has recently been implicated in various forms of cancer in humans, including breast cancer. However, the possible implication of this enzyme in canine mammary cancer has never been addressed. The objective of this proposal is to study the mechanisms involved in the synthesis of prostaglandins in mammary tumors in the dog. We first plan to study the expression of the enzyme COX-2 in benign and malignant mammary tumors in dogs. We then want to characterize the regulation of the enzyme COX-2 in cultures of cells derived from canine mammary tumors. It is expected that a better comprehension of the mechanisms involved in COX-2 expression in canine mammary cancer will help design new strategies for cancer therapy and prevention in the canine species.

Active Grant No. 2254. Heritable and Sporadic Genetic Lesions in Canine Lymphoma and Osteosarcoma

Principal Investigators. Jaime Modiano, VMD, PhD; AMC Cancer Research Center; Matthew Breen, PhD; North Carolina State University

Sponsor(s). Akita Club of America, Inc., American Belgian Tervuren Club, Inc., American Bloodhound Club, American Boxer Charitable Foundation, Atlantic States Briard Club, Inc., Bernese Mountain Dog Club of America, Briard Club of America, Bull Terrier Welfare Foundation, Flat-Coated Retriever Foundation, German Wirehaired Pointer Club of America, Golden Retriever Foundation, Irish Wolfhound Club of America, Inc., Mastiff Club of America, Medallion Rottweiler Club, Nestle Purina PetCare Company, Newfoundland Club of America Charitable Trust, Otterhound Club of America, Portuguese Water Dog Club of America, Inc., Rhodesian Ridgeback Club of the United States

Abstract: Lymphoma (cancer of lymph glands) and osteosarcoma (bone cancer) are two common cancers of dogs with remarkable breed predisposition. Lymphoma accounts for approximately 20 percent of all canine tumors, and > 80 percent of cancers originating from blood cells. Osteosarcoma is the most common bone tumor in dogs, accounting for 85 percent of skeletal cancers. All cancers have a genetic basis, and in effect, these conditions represent various diseases, each sharing one or few genetic abnormalities that contributes to overall risk and treatment response. However, a means does not exist to identify individuals at risk, or tumors that are likely to respond to conventional therapy. We have identified individual genes and larger regions within the genome that appear to be important in canine cancer. For this project, we propose to confirm the frequency and significance of these genetic anomalies in lymphoma and osteosarcoma of Golden Retrievers, Rottweilers, Irish Setters, and Bernese Mountain Dogs This work will begin to determine which of these anomalies may be heritable and which may be sporadic, and pave the way to apply this knowledge for clinical benefits by providing potential

targets for treatment, and tools to define individual risk to develop these types of cancer or produce cancer-prone progeny.

Active Grant No. 2465: Identification and Characterization of Genetic Mutations in Canine Mast Cell Tumors

Principal Investigator: Cheryl London, DVM, PhD; University of California, Davis Sponsor(s). American Boxer Charitable Foundation, American Bullmastiff Association, Chinese Shar-Pei Club of America Charitable Trust, Collie Health Foundation, French Bulldog Club of America, Golden Retriever Foundation, Jeffrey Pepper, Rhodesian Ridgeback Club of the United States, Staffordshire Bull Terrier Club

Abstract: The most common malignant tumor in dogs is the mast cell tumor (MCT, a form of skin cancer), occurring with an incidence of close to 20% in the canine population. MCTs range from relatively benign to extremely aggressive, leading to tumor spread and eventual death. Particular breeds of dog are at risk for the development of this tumor, indicating a role for genetic factors. We have previously identified mutations in the gene c-kit in 30-50% of dog MCTs. c-Kit plays a critical role in regulating the growth and function of normal mast cells, and as the mutations we discovered cause uncontrolled function of c-kit, it is likely they influence MCT development in dogs. This proposal will establish a prospective tumor registry of dog MCTs to be used for investigation of the true incidence of c-kit mutations within specific dog breeds. Moreover, the studies outlined in this grant will identify additional genetic mutations present in dog MCTs that can be used for the development of new, targeted therapeutics. In summary, this work will provide a much more detailed understanding of dog MCTs, thereby building a framework for the development of new therapies and strategies for disease prevention.

Pending Grant No. 249: Genomics of Canine Brain Neoplasia
Principal Investigators: Matthew Breen, PhD and Natasha Olby, PhD; North Carolina State
University

Abstract: Genetic aberrations underlie many different types of cancer. Identification of these aberrations provides important information on the malignancy of different cancers, but until recently it has been extremely labor intensive to screen for such anomalies. Gene expression profiling is a technique that provides information on the level of expression of thousands of genes in a single assay, greatly improving the efficiency of screening for genetic aberrations. We hypothesize that there are tumor-specific differences in gene expression in canine brain tumors, which will correspond to and be predictive of clinical outcome. As certain breeds of dog (e.g. the Boxer, Boston Terrier and Golden Retriever) are predisposed to developing brain tumors, we may find breed specific genetic aberrations that are associated with the development of brain tumors. In this study, we will apply gene expression profiling to tissue taken from naturally occurring brain tumors (obtained during routine diagnosis and treatment) to identify tumor specific differences in gene expression. We will correlate these differences with tumor type and the clinical course of disease to identify prognostic factors for survival. This study may enable us to identify prognostic factors and novel therapeutic targets applicable to many different types of canine cancer.

Pending Grant No. 2629: Clinical and Immunological Outcomes in Dogs with Osteosarcoma Treated with Intratumoral Interleukin-12 Microspheres

Principal Investigator. Stuart Helfand, DVM; University of Wisconsin, Madison Sponsor(s): Akita Club of America, Inc., American Bloodhound Club, American Boxer Charitable Foundation, American Bullmastiff Association, American German Shepherd Dog Charitable Foundation, Borzoi Club of America, Flat-Coated Retriever Foundation, Golden Retriever Foundation, Great Pyrenees Club of Puget Sound, Irish Wolfhound Club of America. Inc., Irish Wolfhound Foundation, Jeffrey Pepper, Newfoundland Club of America Charitable Trust, Rottweiler Health Foundation, St. Bernard Club of America, Starlight Fund Abstract: Appendicular osteosarcoma, or bone cancer of the limbs, is an important tumor in dogs representing nearly 10 percent of all canine cancers. Despite progress in treating canine osteosarcoma using a combination of limb amputation and chemotherapy, life expectancy is not usually extended by more than 6-10 months compared to amputation alone. Death is due to dissemination of cancer cells beyond the leg and it is estimated that the cancer has already spread in at least 95% of dogs when it is initially diagnosed. Any pet owner who has lived through this disease can attest to the heartbreak of the news that tumor masses, initially undetectable at the time of diagnosis, have been discovered on a chest X-ray. Novel treatment regimens are urgently needed to improve the lives of large breed dogs such as Golden, Labrador an other Retriever4s, Rottweilers, Irish Wolfhounds, Great Danes, German Shepherds and others that are at greatest risk for developing this cancer. Stimulating the immune system of dogs with cancer has been a goal of veterinary cancer researchers for more than 20 years and osteosarcoma is a tumor that has shown positive responses to some of these interventions. This research proposes to add a potent new form of immunostimulation to the standard treatment for canine osteosarcoma. This strategy uses a powerful stimulant of the immune system called interleukin-12 (IL-12) that has been shown to induce strong anti-tumor responses in experimental animal models. Stimulated by IL-12, immune cells tolerant of cancer can be triggered to kill cancer cells throughout the body, resulting in the generation of an army of deadly circulating killer immune cells specific for the cancer. What's more, this type of cell has long-term memory for the cancer, in much the same way vaccination results in long-term immunity to infectious disease. Our laboratory has shown that IL-12 enhances killing of osteosarcoma cells by immune cells from dogs with this cancer. Now, we propose that injection of IL-12 directly into limb osteosarcoma using a novel (microsphere) formulation resulting in slow IL-12 release within the tumor environment will promote active anti-tumor immunity in dogs with osteosarcoma and lengthen their survival time. A number of pertinent immunological questions will also be addressed.

Active Grant No 2632 Analysis of Gene Expression in Canine Tumors Principal Investigator Richard W. McCombie, PhD; Cold Spring Harbor Laboratory Abstract: In recent times our understanding of human cancer has changed considerably. We know that a given type of cancer, such as breast cancer, is comprised of a number of subtypes, based on specific genetic changes that occur in a given patient. More and more, these genetic differences will guide not only our understanding of this disease, but the treatment and diagnosis of specific patients. We are proposing to use some of the same tools by which these molecular changes are found in human tumors to study cancer in the dog. By doing this we hope to see how closely canine cancer parallels the disease in humans. In addition to letting us profile the

disease in molecular terms in dogs, it will let us determine to what extent we can use the vast data set from humans to understand the molecular nature of canine cancer.

Active Grant No. 2646: Characterization of Receptor Tyrosine Kinase Dysfunction in Malignant Histocytosis

Principal Investigator: Cheryl London, DVM, PhD; University of California, Davis Sponsor(s) American Boxer Charitable Foundation, Bernese Mountain Dog Club of America, Flat-Coated Retriever Foundation, Golden Retriever Foundation

Abstract: Malignant histiocytosis (MH), while rare in people, occurs frequently in certain breeds of dogs including Rottweilers, Golden Retrievers, Flat-Coated Retrievers and Bernese Mountain Dogs. There is no effective therapy for this disease and nearly all patients die within two to four months of diagnosis. The purpose of this proposal is to evaluate MH tumor specimens for mutations in genes that may contribute to the development of this devastating cancer. The genes of interest are those that code for proteins known as growth factor receptors. These proteins are present on the surface of the cell and when stimulated by growth factors, signal into the cell promoting cell survival and growth. Dysregulation of growth factor receptors is a common mechanism through which normal cells undergo transformation into cancer cells. Significant research has been directed towards the development of inhibitors capable of blocking the function of dysregulated receptors. Recent success of this approach has been realized with the inhibitor Gleevec in the treatment of chronic myelogenous leukemia in people. The purpose of this proposal is to identify growth factor receptors that are dysregulated in MH to provide the foundation for future clinical application of growth factor receptor inhibitors in the treatment of MH.

Pending Grant No. 2657: Role of Circulating Endothelial Cells in Physiologic and Pathologic Angiogenesis

Principal Investigator: Steven W. Dow, DVM, PhD; Colorado State University Sponsor(s) American Boxer Charitable Foundation, Collie Health Foundation, Flat-Coated Retriever Foundation, Golden Retriever Foundation, Great Pyrenees Club of Puget Sound, Saluki Club of America, Inc., Saluki Health Research, Inc., St. Bernard Club of America, Starlight Fund Abstract: Formation of new blood vessels (angiogenesis) is crucial to a number of important processes, including healing wounds, resolving inflammation, and stimulating the growth of tumors. Recent studies indicate that many of the new cells that are used to form new blood vessels actually come from the bloodstream. This observation raises several important questions, including how different disease states may affect that numbers of these cells in circulation and how the numbers of these circulating endothelial cells may be regulated. We propose to evaluate new assays to measure the numbers these circulating endothelial cells in the bloodstream of dogs, using small blood samples. These assays will be used to measure the numbers of these endothelial cells in the bloodstream of dogs with conditions that may affect formation of new blood vessels, including surgery, tumors, and clotting disorders. Circulating endothelial cells will also be grown in tissue culture and characterized. These studies will provide important new information on the role of the bone marrow and circulating endothelial cells in blood vessel formation in dogs and may help in the development of new strategies to either inhibit or stimulate blood vessel formation in different diseases.

Pending Grant No. 2667: Cellular Genomics - Molecular Cytogenetic Investigation of Canine Soft Tissue Sarcomas

Principal Investigator Matthew Breen, PhD; North Carolina State University Sponsor(s) · American Boxer Charitable Foundation, Bernese Mountain Dog Club of America, Collie Health Foundation, Flat-Coated Retriever Foundation, Golden Retriever Foundation Abstract: It has been established that non-random chromosome aberrations are characteristic of specific types of many different human cancers. The knowledge of such aberrations has identified areas of the human genome to be targeted for further research. In the dog the extent and identity of chromosome aberrations associated with specific cancers is still largely unknown. In certain breeds, such as the Flat-Coated Retriever and Bernese Mountain Dog, soft tissue sarcomas account for up to 50 percent of all malignant tumors and thus represent a serious health and welfare issue for those breeds. These tumors are difficult to classify by conventional means and so attention is required to develop alternative approaches. Human soft tissue sarcomas have been demonstrated to be associated with specific chromosomal aberrations that have been shown to have both diagnostic and prognostic significance. This proposal will make use of major recent advances in canine molecular cytogenetics to identify recurrent chromosome aberrations associated with canine soft tissue sarcomas, in particular those of histiocytic origin. This project will identify areas of the canine genome associated with such cancers for further investigation at the sub-chromosomal level.

Pending Grant No. 267. An Investigation into Combined Molecular Approaches to Treat Hemangiosarcoma

Principal Investigator: David J. Argyle, BVMS, PhD; University of Wisconsin, Madison Abstract: Hemangiosarcoma (HSA) is a common and fatal cancer in dogs for which there is no effective treatment. Despite surgery and intensive chemotherapy, the median survival time for dogs diagnosed with HSA is little more than six months. From our own studies on canine cancer, expression of the enzyme telomerase allows cancer cells to become immortal and has emerged as a central and near universal marker of cancer, making it a candidate target for novel therapies. In this study we will explore the value of telomerase inhibition to treat HSA using the novel mechanism of RNA interference (RNAi). Our hypothesis is that potent inhibition using this technique will inhibit the immortal phenotype of the cancer cells and cause them to die. However, it is possible that a combined approach, targeting two molecular pathways, may offer greater therapeutic benefit. Consequently, we will also explore the potential synergism of combining telomerase inhibition with an alternative inhibitor of a further mechanism involved in cancer (receptor tyrosine kinase inhibition) on targeting this highly malignant tumor In this we will use in vitro cell culture techniques for initial inhibition studies followed by studies in a novel canine HSA model system to ascertain the potential clinical merit of this approach for dogs with HSA.

Pending Grant No 272: Oligonucleotide Microarray Gene Expression Profiling of Canine Lymphoma

Principal Investigator William C Kisseberth, DVM, PhD; Ohio State University Abstract: Lymphoma is one of the most common cancers seen in the dog. Current methods of classifying lymphoma neither explain nor predict its variable clinical behavior. While the majority of canine lymphomas appear microscopically similar and affected dogs show similar clinical signs, the clinical course of the disease can vary significantly in patients with

microscopically identical tumors with identical clinical signs. This heterogeneity in behavior is particularly evident with respect to response to chemotherapy. Although the majority of patients initially respond well to chemotherapy, some are disease-free for a few months, while others remain disease-free over two years. Clearly, microscopic and initial clinical appearances inadequately explain the variable clinical behavior. In order to better understand and explain these differences, we will create and develop a specialized dog lymphoma gene microarray; a new tool that can be used to determine which groups of genes are important in different subtypes of lymphoma. Ultimately, by identifying these important groups of genes, we hope to 1) provide better prognostic information regarding individual tumor clinical behavior, 2) identify important groups of genes that characterize unique lymphoma sub-types, and 3) identify new molecules or genes that can be targets for development of new drugs to treat lymphoma.

Pending Grant No. 297: Identification of a Chromosomal Region Associated with the Development of Osteosarcoma in the Scottish Deerhound

Principal Investigator: Marlene Hauck, DVM, PhD; North Carolina State University Abstract: Osteogenic sarcoma (OSA) is the most common malignant bone tumor in dogs. The etiology of canine osteogenic sarcoma (OSA) is largely unknown. Reports have noted an association between prior local radiation therapy, bone infarcts, and prior fractures (fixed with metallic implants) and the development of bony tumors. Veterinarians, given the disproportionate incidence in many breeds, have long suspected a hereditary basis for canine OSA. The Scottish Deerhound is a giant breed of dog with a ten percent estimated incidence of osteosarcoma. By studying family groups of dogs within this breed, we hope to determine both the mechanism of inheritance as well as the region of DNA associated with their high risk of developing OSA. Determination of the genetic regions (or genes) responsible for the high incidence of OSA in breeds such as the Scottish Deerhound will allow further understanding of the mechanisms underlying neoplastic transformation, enable genetic testing for carriers across all breeds, as well as present new therapeutic targets.

Deafness

Active Grant No 1870: Genetics of Hereditary Deafness in the Domestic Dog Principal Investigator: Keith E. Murphy, PhD; Texas A&M University Sponsor(s): Dalmatian Club of America Foundation, Inc., English Setter Association of America

Abstract: Congenital deafness has been reported for approximately 60 breeds, and can potentially appear in any breed. The disorder is usually associated with pigmentation patterns, where increasing amounts of white in the hair coat increase the likelihood of deafness. Pigment-associated inheritance of deafness is not restricted to dogs. Similar defects have been reported for mice, mink, pigs, horses, cattle, cats and humans. Waardenburg syndrome type 2 (WS2) in humans presents with congenital deafness and hypopigmentation of the eyes and head hair. This is an autosomal dominant disorder with incomplete penetrance, meaning that individuals that inherit the disorder may not show all of the components of the syndrome. Mutations in the mitf gene cause at least twenty percent of cases of WS2. Piebaldism in humans most commonly presents with congenital patches of white skin and hair-lacking melanocytes on certain parts of the body, although deafness is a characteristic of the disorder in an unknown percentage of

affected individuals. Mutations in the kit gene are responsible for typical cases of piebaldisms. We believe that deafness is dogs may result from mutations in the canine mitf/kit genes. Examination of these genes and identification of causative mutations will aid in understanding the etiology of deafness.

Active Grant No. 2264: Whole Genome Screens Using Microsatellite Markers in Genetic Analyses of Hereditary Deafness in the Dalmatian

Principal Investigator: Keith E. Murphy, PhD; Texas A&M University

Sponsor(s) Dalmatian Club of America Foundation, Inc.

Abstract: Hereditary deafness has been reported for approximately 60 breeds, and can potentially appear in any breed. The disorder is often associated with pigmentation patterns, with increasing amounts of white in the hair coat increasing the likelihood of deafness. Pigment-associated inheritance of deafness is not restricted to dogs. Similar defects have been reported for mouse, mink, pig, horse, cattle, cat and human. High incidences of deafness are found in the Dalmatian and English Setter. Deafness in these breeds presents unilaterally (with no preference for either ear) or bilaterally and recent research suggests that more than one gene is involved. Therefore, in an effort to identify such genes, we will carry out analysis of the entire genome rather than restricting studies to one or two genes. By using this approach, termed linkage analysis, we hope to identify markers present in affected dogs. Informative markers will allow selection and subsequent examination of candidate genes for mutations that play roles in deafness of the aforementioned breeds. The long-term objectives of this work are to 1) develop marker or genebased tests to identify carriers and to reduce the incidence of deafness, and 2) understand the etiology of deafness in the Dalmatian and English Setter.

Epilepsy

Pending Grant No. 224: Characterizing Idiopathic Epilepsy in the Poodle, Giant Schnauzer, English Mastiff: An Assessment of Inheritance

Principal Investigator. Anita M. Oberbauer, PhD; University of California, Davis Abstract: Idiopathic epilepsy afflicts many dogs, in fact, 35 Parent Clubs list epilepsy as being a major health concern. In some breeds, epilepsy has been established to be an inherited disorder although the mode of inheritance differs. For the Poodle, Giant Schnauzer, and English Mastiff, three breeds in which epilepsy is a recognized health concern, the mode of inheritance, heritability and relative incidence have not been defined. Collection of health information, pedigrees and DNA samples will provide the necessary data needed to characterize this disorder in these breeds. Survey data describing the characteristics of the seizures (frequency, age of onset, and intensity) will be collected and statistically analyzed to generate reliable heritability estimates of seizuring. Once we have determined how heritable epilepsy is within these breeds, more advanced statistical procedures will be employed to ascertain whether epilepsy is a complex disorder involving many genes or whether a single gene regulates the seizures. If the statistical analyses suggest a major gene affects the seizures within each breed, then it becomes possible to identify a genetic marker linked to that gene. Determining heritability and mode of inheritance of epilepsy in the Poodle, Giant Schnauzer, and English Mastiff will prove valuable for breeders to make informed, health-based breeding decisions.

Active Grant No. 2252. Canine Epilepsy: Determining the Mode of Inheritance, Mapping the Genes, and Developing a Linkage Test

Principal Investigator: James R. Mickelson, PhD and Ned Patterson, DVM; University of Minnesota

Sponsor(s): English Springer Spaniel Field Trial Association Foundation, National Beagle Club, Vizsla Club of America, Vizsla Club of America Welfare Foundation

Abstract: We propose to continue our molecular genetic studies to develop a screening linkage test for predicting epilepsy in Beagles, English Springer Spaniels and Vizslas. Preliminary results of our genetic marker studies in these breeds indicate that we will be able to find linked markers and the chromosomal segment containing the epilepsy gene given sufficiently large and informative pedigrees. The onset of seizures in dogs with epilepsy is typically from one to five years of age. The late onset means that often a dog has already been bred before it is know to be affected. In some individuals, seizures are well controlled with anticonvulsant medications, but a significant number of dogs have "refractory" seizures needing high doses of medications to achieve control. The severity of seizures may be such that the owner elects to have the dog euthanized. A genetic test for epilepsy would allow breeders to screen potential breeding animals for this common, frustrating, and potentially devastating disorder prior to making breeding decisions. Our genome mapping approach to identifying the region of the canine genome containing the defective gene will ultimately lead to the prediction of candidate genes that can be characterized to define the precise defect responsible.

Active Grant No. 2304: Continued Investigation into the Molecular Genetic Causes of Canine Epilepsies

Principal Investigator: Gary S. Johnson, DVM, PhD; University of Missouri Sponsor(s): American Spaniel Club Health Foundation, AWS Partners, Collie Health Foundation, Dalmatian Club of America Foundation, Inc., English Springer Spaniel Field Trial Association Foundation, Greater Swiss Mountain Dog Club of America, Inc., Irish Setter Club of America Foundation, Irish Water Spaniel Club of America, St. Bernard Club of America, Standard Schnauzer Club of America, Welsh Springer Spaniel Club of America Abstract: In our on-going study of canine epilepsy, we have found that most canine epilepsy families do not follow the simple inheritance patterns described by Gregor Mendel. This suggests that epilepsy results when a dog inherits mutations in two or more different genes. Mutations involving two or more genes are also thought to be responsible for the vast majority of human epilepsy that occurs in families. Mendelian inheritance is encountered in rare types of human epilepsy and in many of these families the epilepsy-causing mutation has been found by genome mapping. We appear to have found Mendelian inheritance in at least one canine family and if this holds true we will attempt to map the mutation. One unexpected finding that is not seen in the human epilepsy families is the predominance of males among the affected dogs in several dog breeds. A similar inheritance pattern in human disease has been attributed to paired mutations on the X chromosome and in the mitochondrial DNA in these breeds. Our ultimate goal is to produce a DNA marker that breeders can use to avoid matings that will result in new generations of epileptic dogs.

Active Grant No. 2614 Refining the Genetic Linkage for Idiopathic Epilepsy in the Belgian Tervuren and Sheepdog

Principal Investigator: Anita M. Oberbauer, PhD; University of California, Davis Sponsor(s): American Belgian Tervuren Club, Inc., Belgian Sheepdog Club of America Abstract: Idiopathic epilepsy affects over 30 different AKC dog breeds and was identified as the top canine health issue in the 2001 AKC/CHF Parent Club Survey. Preliminary studies in the Belgian Tervuren and Sheepdog indicate that although this is a heritable disorder governed by many genes, there exists a single gene of very large effect on the incidence of epilepsy. The epileptic condition influenced by this particular gene is inherited as an autosomal recessive. Preliminary data in our laboratory, derived from screening the DNA of affected dogs, identified a chromosomal region appearing to be associated with epilepsy. The specific objective of this project is to refine this initial association to a very discrete region of the DNA in order to ultimately develop a genetic marker associated with epilepsy. Such a genetic marker will permit the identification of carriers of epilepsy in Belgian Tervuren and Belgian Sheepdogs, and possibly other breeds, prior to breeding with the end result that breeders can make informed, health-based breeding decisions.

Pending Grant No. 266 Canine Epilepsy: Mapping the Genes and Developing a Linkage Test (Continuing Studies)

Principal Investigator James R. Mickelson, PhD and Ned Patterson, DVM; University of Minnesota

Abstract: We seek to continue our molecular genetic studies to develop a screening linkage test for predicting epilepsy in Beagles, English Springer Spaniels, and Vizslas and to institute new studies in Welsh Springer Spaniels, Greater Swiss Mountain Dogs, Irish Setters, Otterhounds, Samoyeds and Keeshonden. Preliminary results of our genetic marker studies indicate that we will be able to find linked markers and the chromosomal segment containing the epilepsy gene given sufficiently large and informative pedigrees. The late age of onset of seizures in dogs with epilepsy means that a dog has often already been bred before it is diagnosed as affected. In some individuals seizures are well controlled with anticonvulsant medications. However, a significant number of dogs have "refractory" seizures needing high doses of medications to achieve control or the severity of seizures may be such that the owner elects to have the dog euthanized. Our genome mapping approach to identifying the regions of the canine genome containing the defective genes will ultimately lead to genetic tests for epilepsy that would allow breeders to screen potential breeding animals for this common, frustrating, and potentially devastating disorder.

Eye Disease (Other than PRA)

Active Grant No. 2291 Molecular Genetic Causes for Canine Lens Luxation and Glaucoma Principal Investigator Gary S. Johnson, DVM, PhD; University of Missouri Sponsor(s) American Sealyham Terrier Club, Basset Hound Club of America, Inc., Bouvier Health Foundation, Bull Terrier Welfare Foundation, Dandie Dinmont Terrier Club of America, Inc., Dandie Dinmont Trust Fund, Health & Rescue Foundation of the Petit Basset Griffon Vendeen Club of America, Jack Russell Terrier Research Foundation, Miniature Bull Terrier

Club of America, Montgomery County Kennel Club, Inc., Tibetan Terrier Club of America, Welsh Terrier Club of America, Inc.

Abstract: Heritable lens luxation, if not treated promptly, will induce secondary glaucoma. In addition, heritable primary glaucoma can cause secondary lens luxation. Since it is not always known whether lens luxation or glaucoma is the primary disease, we believe it is rational to study both diseases together. One or the other of these diseases is responsible for loss of sight in the Basset Hound, Border Collie, Dandie Dinmont Terrier, Jack Russell Terrier, Miniature Bull Terrier, Petit Basset Griffon Vendeen, Sealyham Terrier, Tibetan Terrier, Welsh Springer Spaniel, and Welsh Terrier. We are attempting to produce DNA marker assays that will identify dogs with the mutant gene responsible for lens luxation and glaucoma. Early identification of these dogs would enable dog owners and their veterinarians to instigate measures to preserve their dogs' sight and to adjust breeding practices to minimize or eradicate the disease in their breeds.

Heart Disease

Active Grant No 2002: Evaluation of the Clinical Outcome of Asymptomatic Adult Boxers with Ventricular Arrhythmias Over a Four-Year Period

Principal Investigator: Kathryn Meurs, DVM, PhD; Ohio State University

Sponsor(s): American Boxer Charitable Foundation

Abstract: Heart disease in the Boxer was initially documented in the 1980s and referred to as a Boxer cardiomyopathy. More recent studies have confirmed that this disease is inherited and is primarily characterized by disturbances in the cardiac electrical system, fainting episodes and sudden death. The inherited nature of the disease has led to increased interest in the screening of dogs for the disease by electrocardiogram, ultrasound and Holter monitoring prior to using them for breeding. However, the interpretation of the results is difficult, since many adult Boxers have some abnormalities detected on at least one of the tests and there is no available information regarding the relationship between these findings and the likelihood of development of clinical signs. The objective of this study is to evaluate the correlation between specific cardiac parameters (Ventricular Premature Contractions (VPC) number, grade of arrhythmia, heart rate, etc.) detected by electrocardiogram, ultrasound, and Holter monitoring and the development of clinical signs including fainting, sudden death and congestive heart failure in 130 adult Boxers previously diagnosed with heart disease.

Active Grant No 2250: Diet-Related Taurine Deficiency in Newfoundland Dogs and Associated Cardiac Insufficiency

Principal Investigator Robert Backus, DVM, PhD; University of California, Davis

Sponsor(s): Newfoundland Club of America Charitable Trust

Abstract: Dilated cardiomyopathy (DCM) is the most common acquired cardiovascular disease of dogs. Of suggested nutritional causes, taurine deficiency appears most evinced. But in dogs, such a need for dietary taurine is not generally recognized. Dogs are known to have the metabolic capacity to synthesize taurine from the dietary sulfur amino acids, cysteine and methionine (Hayes 1988). Diet-related taurine deficiencies and associated dilated cardiomyopathies have been reported in large breed dogs (Torres et al. 2000). The present investigators (Backus et al. 2000) have recently reported taurine deficiency (52 percent) and

cardiac insufficiency (10 percent) among a group of 21 privately owned Newfoundland dogs. The proposed research will estimate the prevalence of a possible unrecognized, widespread, taurine deficiency in the Newfoundland breed. This will involve conducting non-invasive tests in a clinical setting. Dogs will be evaluated for heart and retinal disease. In selected normal and taurine deficient dogs, blood and urine will be collected to determine taurine and sulfur amino acid concentrations. Feeding trials will evaluate the effect of diet composition on taurine concentrations and excretion. A possible genetic contribution to taurine deficiencies will be assessed by pedigree analyses.

Active Grant No. 2266: Linkage Analysis of Familial Dilated Cardiomyopathy in the Doberman Pinscher

Principal Investigator. Kathryn Meurs, DVM, PhD; Ohio State University Sponsor(s) Doberman Pinscher Club of America

Abstract: Dilated cardiomyopathy (DCM), a primary heart muscle disorder characterized by poor cardiac function, is inherited in the Doberman Pinscher. Therapy for DCM does not cure or even successfully control the clinical signs. The inability to control the disease has led to increased interest in disease prevention. Prevention will likely be achieved through careful selection of unaffected dogs for breeding. However, since DCM is often not apparent until later in the adult life of the dog, many dogs are selected for breeding before they are found to be affected. A blood test that could identify affected animals before they are used for breeding would greatly decrease the prevalence of DCM. The study proposed here is a continuation of a study funded by the AKC/CHHHF that recruited and screened families of Doberman Pinschers with DCM. Pedigrees and DNA samples have been collected. Genetic analysis is now being performed on three families of Doberman Pinschers with DCM. Genetic markers will be evaluated to determine if they are associated with the development of DCM in the Doberman Pinscher. The identification of a genetic marker linked to DCM will be the first step in the development of a screening blood test.

Active Grant No 2267. Linkage Analysis of Familial Subvalvular Aortic Stenosis in the Newfoundland Dog

Principal Investigator: Kathryn Meurs, DVM, PhD; Ohio State University Sponsor(s) Newfoundland Club of America Charitable Trust, NewPenDel Newfoundland Club Abstract: Subvalvular Aortic Stenosis (SAS) is a familial congenital heart disease characterized by a fibrous ridge located below the aortic valve. Affected dogs are at risk of developing an infection on the aortic valve, congestive heart failure or sudden death. This is a familial defect in the Newfoundland dog; therefore affected dogs should not be bred. However, mildly affected dogs can be difficult to diagnose without Doppler echocardiography, an expensive test with limited availability. Therefore, there is significant interest in developing a genetic test to screen for SAS. The study proposed builds on a previous study funded by the AKC/CHF to recruit and screen families of Newfoundland dogs with SAS. Dogs have been classified as affected, unaffected but related to affected, or unaffected by Doppler echocardiography. Pedigrees and DNA samples have been collected on several small families. Additional recruitment of grandparents and siblings will complete these families. Genetic analysis will be performed using canine markers and DNA samples from these families. Genetic markers will be evaluated to determine if they are associated with SAS in the Newfoundland. The identification of a genetic marker linked to SAS will be the first step in the development of a DNA screening test.

Active Grant No. 2303: Molecular Analysis of Familial Ventricular Arrhythmias in the Boxer Dog

Principal Investigator: Kathryn Meurs, DVM, PhD; Ohio State University

Sponsor(s): American Boxer Charitable Foundation

Abstract: A heart muscle disease in the Boxer dog was initially documented in the 1980s and referred to as Boxer cardiomyopathy. Studies have confirmed that this is an inherited disease primarily characterized by an electrical disturbance in the heart that may lead to collapsing episodes and sudden death. There is increasing demand for a screening test that could be used to evaluate dogs for the disease before they are selected for breeding purposes. Unfortunately, many of the clinical abnormalities do not become apparent until the dog is several years old. Therefore, there is significant interest in developing a DNA test that could be performed before a dog is selected for breeding. The study proposed here is a continuation of a study in which 268 Boxers were evaluated by physical examination, echocardiography and ambulatory electrocardiography. Dogs have been classified as affected, equivocally affected or unaffected; pedigrees and DNA samples have been collected. Three-generation families have been identified. Linkage and candidate gene analysis will now be performed using canine markers and these DNA samples. The identification of a genetic marker linked to familial ventricular arrhythmias will be the first stem pin the development of a DNA screening test.

Active Grant No. 2306: Evaluation of the Geographical Variation in Familial Nature and Inheritance Patterns of Boxer Dogs with Familial Ventricular Arrhythmias Principal Investigator: Matthew Miller, DVM; Texas A&M University Sponsor(s): American Boxer Charitable Foundation

Abstract: Heart disease in the Boxer dog was initially documented in the 1980s and referred to as Boxer cardiomyopathy. Recent studies have confirmed that this disease is inherited and is primarily characterized by disturbances in the cardiac electrical system, fainting episodes and sudden death. The inherited nature of the disease has led to increased interest in the development of a genetic test for the disease that could be used to diagnose affected dogs before they were used for breeding. However before investing significant time and resources into the development of a genetic test it is imperative that the familial nature of the disease be well understood. To this date, both studies suggesting a familial nature to Boxer ventricular arrhythmias have been performed in eastern regions of the United States. It is important to perform similar genetic evaluations in a unique geographical region to confirm that the familial nature and the mode of inheritance are constant. The study proposed here will provide an increased understanding of the pattern of inheritance and genetic etiology of this disease within the United States as well as to provide a bank of DNA samples from affected dogs.

Active Grant No. 2429: The Assessment of Ejection Murmurs in the Boxer Dog Principal Investigator: Kathryn Meurs, DVM, PhD; Ohio State University Sponsor(s). American Boxer Charitable Foundation

Abstract: Subvalvular aortic stenosis (SAS) is a common, inherited birth defect of the heart. SAS often affects Boxers and impacts breeding programs. Severely affected dogs are at risk for heart failure, heart infection, and sudden death. Veterinarians usually identify SAS by listening for a heart murmur. In over 50 percent of Boxers, a murmur compatible with SAS is found, prompting sophisticated ultrasound imaging (echocardiography) and blood flow studies (Doppler). Even these tests may not distinguish a stressed or excited, but otherwise normal dog,

from one with mild SAS. This uncertainty is a source of frustration to Boxer breeders. The proposed study explores causes of soft murmurs and increased blood velocities in Boxers. Extensive noninvasive ultrasound studies comparing affected and unaffected dogs are proposed. Furthermore, the origin of these soft murmurs is investigated in a subgroup of Boxers. In these clinical evaluations we will employ "gold standard" methods of X-ray contrast angiography (die), direct (catheter) measurement of blood flow in the heart, and recording of heart murmurs within the heart and blood vessels. We hope to answer the pivotal questions: are these soft murmurs and increased blood velocities really due to SAS, or do they simply represent a normal physiologic event?

Active Grant No 2620: Determination of the Clinical Phenotype and Inherited Nature of Familial Subvalvular Aortic Stenosis in the Rottweiler Dog Principal Investigator: Kathryn Meurs, DVM, PhD; Ohio State University Sponsor(s): Medallion Rottweiler Club, Rottweiler Health Foundation Abstract: Subvalvular aortic stenosis (SAS) is a congenital heart disease characterized by a fibrous ridge located below the aortic valve. Affected dogs are at risk of developing heart valve infections, congestive heart failure or sudden death. This trend has been reported with increasing frequency in the Rottweiler. The defect has been shown to be inherited in the Newfoundland breed, however the inherited nature of the disease in other breeds of dogs, including the Rottweiler, is unknown. The objectives of this study are to define the clinical presentation of SAS in the Rottweiler, compile pedigrees with reference to defined clinical cardiovascular status and evaluate for specific modes of inheritance, and accumulate a databank of clinical information, pedigrees and DNA from Rottweilers affected with SAS and all surviving family members. This study will help define both the clinical attributes and inherited nature of the disease. Information obtained in this study will provide the background for developing both screening and treatment programs and provide the initial materials for molecular studies to be performed in the future.

Hip Dysplasia

Pending Grant No 212. Development of a New Resource for Positional Cloning of Hip Dysplasia Genes: A High Density SNP Map of Canine Chromosome One Principal Investigator. Elaine Ostrander, PhD; Fred Hutchinson Cancer Research Center and Francis Galibert, PhD; CNRS, France

Abstract: Genome maps are essential for identifying disease genes. The current canine map is composed of several thousand markers, and as a result, has proven useful for localizing several disease genes. A much more highly refined map is necessary if we are to actually clone disease genes of interest (not just identify their location) and, subsequently, develop appropriate genetic tests. This proposal aims at developing the technology to do that, focusing on a test case on chromosome one, where two genes associated with hip dysplasia in the Portuguese Water Dog have been mapped. The goal is to determine the location and linear order of many hundred small variants called "SNPs" throughout chromosome one. The resulting SNP map can then be used 1) by researchers on this project to identify the culprit gene(s) in PWD and 2) by anyone studying hip dysplasia in any breed of dog to determine if the same chromosomal region that is mutated in the Portuguese Water Dog is similarly responsible for disease in other breeds.

Joint Trauma

Active Grant No. 2405. Inhibition of Collagenolysis in Canine Cranial Cruciate Ligament During Rupture

Principal Investigator: Peter Muir, DVM; University of Wisconsin, Madison Sponsor(s): Great Pyrenees Club of America, National Amateur Retriever Club, Orthopedic Foundation for Animals

Abstract: The long-range goal of this work is to study tissue repair in the canine cruciate ligament and to determine what causes cruciate rupture. Our objective is to determine whether the presence of two enzymes, tartrate-resistant acid phosphatase and cathepsin K, causes excessive breakdown of collagen within the cruciate ligament, and weakening of the ligament. It is currently believed that excessive expression of this type of enzyme is an important factor causing cruciate rupture. Having determined whether these enzymes cause breakdown of collagen within the cruciate ligament, this knowledge will offer new insight into cruciate rupture in dogs and facilitate development of new treatments. To accomplish the objective of this application, we will determine whether these enzymes are released into the knee joint fluid in dogs with cruciate rupture. Using a tissue culture technique with pieces of ruptured ligament collected from surgical patients, we will also determine whether treatment of the ligament with enzyme inhibitors prevents excessive breakdown of ligament collagen. Upon completion of this work, we expect to have gained a detailed understanding of the role of these ligament-dissolving enzymes in cruciate rupture. We also expect that the results will lead to development of new medical treatments for cruciate rupture.

Pending Grant No. 247: The Study of the Genetics of Cranial Cruciate Ligament Disease in the Dog

Principal Investigator: Max Rothschild, PhD, Iowa State University

Abstract: Cranial cruciate ligament disease (CCLD) is the cause of limping in nearly 20 percent of all dogs that are taken by their owners to veterinarians for lameness. CCLD causes instability in the knee, swelling and pain. Surgery to stabilize a torn CCL costs individual dog owners thousands of dollars and the dog owning public tens of millions of dollars each year and even with surgery, debilitating arthritis occurs and progresses. CCLD commonly occurs in particular breeds of dogs (e.g. Labrador Retriever, Rottweiler, and Newfoundland) while other breeds (e.g. Greyhound, German Shepherd) rarely develop this problem. When a disease is seen with increased frequency in particular breeds of dogs, this supports the probability that a disease, in this case CCLD, is hereditary. Using this information we have already collected pedigrees and genetic material (DNA) from normal and CCLD affected Newfoundlands. The pedigrees will be used to determine a pattern of inheritance (i.e. simple recessive, sex-linked) for CCLD. The DNA will be used to search for genetic markers that differ between normal and affected dogs and may serve as disease predictors. These markers will then allow identification of carriers of CCLD, and potentially reduction or elimination of CCLD from the dog population.

Kidney Disease

Active Grant No. 2020: Inheritance of Urinary Stone Formation in the Dalmatian Principal Investigator. Danika Bannasch, DVM, PhD; University of California, Davis Abstract: Dalmatians are unique among dogs since they excrete uric acid in their urine rather than allantoin like other dogs. Some male Dalmatians form bladder stones composed of urate. The stones can cause a blockage of the urethra preventing the dog from urinating, which is a lifethreatening problem. After surgical removal of the urinary stones, medical treatment is available to prevent their reoccurrence. The treatment for Dalmatians that form stones requires lifelong medication and frequent visits to the veterinarian for urine testing. All Dalmatians regardless of line or the sex of the dogs produce uric acid in their urine, however only a subset of male Dalmatians has the stone formation problem. The goal of this project is to determine if there is an inherited component to the formation of urate stones in Dalmatians and determine the gene involved.

Active Grant No. 2219: Longitudinal Clinical Study, Mode of Inheritance and Therapeutic Trial of Protein-Losing Enteropathy and Nephropathy in Soft Coated Wheaten Terriers Principal Investigator: Shelly Vaden, DVM, PhD; North Carolina State University Sponsor(s). Soft Coated Wheaten Terrier Endowment Fund Abstract: Soft Coated Wheaten Terriers (SCWT) are at risk for protein wasting diseases of the kidneys and intestines. Clinical signs vary but can be severe, even fatal. We demonstrated that the disease is associated with food allergies. We propose to continue the evaluation of our colony of dogs that are genetically predisposed to this disorder. Some of these SCWT already have the disease. Longitudinal evaluation of these dogs will allow us to characterize the early clinical signs, progressive nature, and mode of inheritance of this disease. These findings will allow for future development of a genetic marker and selective breeding programs aimed at elimination of this disorder from the SCWT population. Once dogs become overtly affected with this disease, we will enter them into a treatment trial, using methods known to be effective in food allergies. Information obtained from this trial will be invaluable in developing treatment regimens for affected SCWT in the general population. The youngest dogs in our colony have been weaned directly to a unique diet to determine if diet can be used to prevent disease. Results from these studies have applicability to other breeds of dogs that are at risk for food allergies and/or specific gastrointestinal and renal diseases.

Active Grant No 2279 Longitudinal Field Studies of Families of Soft Coated Wheaten Terriers Affected with Protein-Losing Enteropathy and/or Protein-Losing Nephropathy and the Foundation of a DNA Bank

Principal Investigator. Meryl Littman, VMD; University of Pennsylvania Sponsor(s) Soft Coated Wheaten Terrier Club of America, Inc.

Abstract: An inherited predisposition for diseases causing protein loss from the intestine (protein-losing enteropathy, PLE) and kidney (protein-losing nephropathy, PLN) has been found in Soft Coated Wheaten Terriers. Dogs often show no signs of illness until middle age, and by then many dogs have been bred. Tissue biopsies commonly show inflammatory bowel disease and immune-mediated glomerulonephritis. The mode of inheritance is not proven, and an environmental trigger may be necessary for expression. After diagnosis, most dogs succumb to their disease within a year despite therapy, and some die suddenly. Currently there is no

predictive test to determine which animals may later become ill (affected), which may be passing on at-risk genes (carriers), and which animals are normal. By studying families of affected dogs and monitoring individuals annually with screening tests to detect early signs of abnormalities, we will study the significance of those changes, the clinical course of these diseases and we will attempt to alter the course with diet changes and medications. We will be able to store and begin to study DNA from affected animals, their families, and geriatric healthy animals, in an effort to find a genetic test to help identify affected, at-risk, and normal individuals.

Active Grant No. 2458: Molecular Genetic Characterization of Canine Cystinuria for the Development of Carrier Tests

Principal Investigator: Paula S. Henthorn, PhD; University of Pennsylvania Sponsor(s): American Bullmastiff Association, Bulldog Club of America Charitable Health Fund, Inc., Mastiff Club of America, Scottish Deerhound Club of America Abstract: Cystinuria is an inherited disease that has been long recognized in dogs and has been documented in nearly 70 breeds. Our previous studies made significant progress with a form of cystinuria that is clinically manifested at an early age such that we can go from seeing an affected animal in the clinic to a DNA-based test in a matter of weeks. We now have access to sufficient pedigrees to make similar progress for cystinuria that presents at a later age, and probably represents a more common form of the disease. Genetic testing is even more important for a disease that has a later age of onset.

Active Grant No. 2462. Molecular Genetic Study of Fanconi Syndrome in Basenjis Principal Investigator Gary S. Johnson, DVM, PhD; University of Missouri Sponsor(s): Basenji Club of America, Inc. & Basenji Health Endowment Abstract: The Basenji Club of America considers Fanconi Syndrome the problem of highest priority in the 2001 Parent Club Survey. Our long-term goals are to identify the mutation responsible for Fanconi Syndrome in Basenjis and to use this information to design a DNAbased test for carriers of this disease. This test would enable Basenji breeders to avoid producing future generations of Basenjis with Fanconi Syndrome. Our specific objectives are (1) to isolate DNA from Basenjis with Fanconi Syndrome and their close relatives; (2) to evaluate variant sequences in mitochondrial DNA from an affected Basenji as possible causes of Fanconi Syndrome; (3) to evaluate a region of canine chromosome 30 that corresponds to a chromosomal region responsible for human Fanconi Syndrome locus and, if necessary, to extend the evaluation to cover the entire chromosome; and (4) to establish an internet website for recruiting DNA samples and disseminating information about Fanconi Syndrome and its management. The proposed experiments should localize the Basenji Syndrome locus if it is in the mitochondrial genome or on canine chromosome 30. If the Fanconi locus is elsewhere in the nuclear genome, the proposed experiments should test the feasibility of finding it by whole-genome mapping in future studies.

Neurological Disease

Active Grant No. 2220T. Characterization of Degenerative Myelopathy in the Pembroke Welsh Corgi

Principal Investigator: Joan Coates, DVM, MS, DACVIM; University of Missouri

Sponsor(s): Pembroke Welsh Corgi Club of America

Abstract: Degenerative myelopathy (DM) is a degenerative spinal cord disease affecting, primarily, but not exclusively older German Shepherd Dogs. Recently, veterinarians have documented DM in the Pembroke Welsh Corgi. Affected dogs show progressive rear limb weakness and eventually paralysis. We suspect that an imbalance of methionine or related substances in the spinal fluid may affect the production of myelin, the fatty covering of the nerves that is essential to their function. Oxidative stress and excitotoxicity also may play a concomitant role in spinal cord degeneration and impaired methionine metabolism. Testing spinal fluid for concentrations of metabolites related to methionine metabolism and oxidative stress may allow dogs at risk to receive treatment before clinical signs of disease become apparent and to be identified before being used for breeding purposes. Investigators will collaborate with the Pembroke Welsh Corgi Club of America by developing an epidemiologic survey to determine the prevalence of DM. DNA will be banked and pedigree data collected from affected dogs and family members while maintaining anonymity so that it will be available for future gene mapping studies. We anticipate that the results of the proposed studies will facilitate accurate diagnosis and monitoring of DM.

Active Grant No. 2232: Cricopharyngeal Dysphagia in the Golden Retriever Principal Investigator: Margret Casal, PhD; University of Pennsylvania Sponsor(s): Golden Retriever Foundation

Abstract: Cricopharyngeal Dysphagia is a swallowing disorder in dogs that is apparent at the time of weaning. It may result in failure to thrive, regurgitation, nasal discharge, pneumonia or even death resulting from recurrent pneumonia. Cricopharyngeal dysphagia has been described in several dog breeds and is suspected to be an autosomal recessive disease in these breeds. It has come to our attention that the disease has been recently described in Golden Retrievers, where the disease is suspected to be a dominant trait with variable expression. The objective of this study is to determine the mode of inheritance and to collect DNA samples to develop a genetic test in the future. To determine the mode of inheritance, careful diagnosis of the presence of absence of cricopharyngeal dysphagia using fluoroscopy is required, as there may be some dogs with a very mild, almost undetectable swallowing disorder. The ultimate goal is to eradicate this devastating disease from the Golden Retriever population.

Active Grant No 2636 Characterization of Cerebellar Cortical Degeneration in American Staffordshire Terriers

Principal Investigator. Natasha Olby, PhD; North Carolina State University

Sponsor(s). Staffordshire Terrier Club of America

Abstract: An inherited degenerative disease affecting the cerebellum has recently been recognized in adult American Staffordshire Terriers. The cerebellum controls coordination of movement, and as a result of degeneration of cerebellar neurons, affected dogs have progressive difficulty with coordination and balance. Signs of the disease start from two to six years of age and are subtle at first, but progress at varying rates until the dog is unable to walk. The late onset

of signs means that dogs can be bred widely before the disease manifests itself, and development of a diagnostic test that can be used to screen dogs for the disease is therefore extremely important. The aims of this study are to describe in detail the clinical and pathological characteristics of this disease, to determine its mode of inheritance, to bank DNA from affected and unaffected dogs for future use, and to identify possible candidate genes for the disease. Affected dogs will be recruited by disseminating information in the disease to owners and breeders of American Staffordshire Terriers with the help of the American Staffordshire Terrier Club. The clinical information and DNA collected in this study will form the basis for future work on identifying the mutation causing this disease. Our ultimate aim is to develop a diagnostic test for cerebellar cortical degeneration in American Staffordshire Terriers.

Pending Grant No. 323: Mapping the Gene Responsible for Hypomyelination in Weimaraner Dogs

Principal Investigator: Ian Duncan, PhD; University of Wisconsin, Madison Abstract: The aim of this project is to map the gene responsible for a central nervous system (CNS) disorder in the Weimaraner breed. This disorder, which appears to be inherited as a single gene defect in an autosomal recessive trait, is widespread throughout North America. Affected dogs develop a severe tremor at ten to fourteen days with consequent high morbidity. Many pups show difficulty in ambulating and nursing, requiring intensive care. Although the tremor in many pups will lessen with time, some are euthanized due to the severity of signs at the time of diagnosis. Affected pups are also not suitable for breeding or the show ring. An identical disorder is also seen in Chow Chows. The long-term goal is to provide a screening test for heterozygotes and find the gene, thus helping to lower the prevalence of the disease. To begin with, we will concentrate on the Weimaraner as we have extensive pedigrees and DNA for this breed. Linkage analysis with microsatellite markers will be applied to map the location of the gene. Then, the linked marker(s) could serve as an indicator of the existence of the disease related allele in the family.

Orthopedic Disease (Other than Hip Dysplasia)

Active Grant No 2460: Molecular Genetic Characterization of Mucopolysaccharidosis Type VI for the Development of a Carrier Test and Relationship to Legg-Calve-Perthes Disease Principal Investigator: Urs Giger; University of Pennsylvania Sponsor(s): Helen Chrysler Greene, Miniature Pinscher Club of America, Inc., Westie Foundation of America, Inc.

Abstract: Legg-Calve-Perthes disease is an orthopedic abnormality causing pain and decreased hip motion. The cause of this disease is unknown, but it is likely to be multi-factorial. It is common in smaller breeds of dog, including the Miniature Pinscher. Similarly, mucopolysaccharidosis type VI (MPS VI) is a disease that causes hip and patellar abnormalities It, too, has been recognized as occurring with increased frequency in the Miniature Pinscher and some other breeds such as Miniature Schnauzers, Welsh Corgis, and Chesapeake Bay Retrievers. It is an autosomal recessively inherited disease caused by a deficiency in the enzyme arylsulfatase B. When the normal canine arylsulfatase gene sequence is know, the mutation responsible for MPS VI in the Miniature Pinscher as well as other breeds can be determined and it will be possible to develop genetic tests. Since MPS VI and Legg-Calve-Perthes disease have

both been diagnosed in Miniature Pinschers and both cause skeletal defects, we hypothesize that at least a subset, if not all, Miniature Pinschers with Legg-Calve-Perthes disease in fact have MPS VI. We propose to develop a genetic test for MPS VI in the Miniature Pinscher and other breeds and to identify individuals with Legg-Calve-Perthes disease to see if they test as normal, carrier or affected in the MPS VI test.

Pending Grant No. 2616. Molecular Analysis of Contributory Factors of Osteoarthritis in Canine Hip Dysplasia

Principal Investigator: Alpana Ray, PhD; University of Missouri Sponsor(s) American Bloodhound Club, Basenji Club of America, Inc. & Basenji Health Endowment, Golden Retriever Foundation, Norwegian Elkhound Association of America, Inc., Random Individuals, Rottweiler Health Foundation, St. Bernard Club of America Abstract: Hip Dysplasia is a common disease of dogs that will ultimately lead to osteoarthritis (OA), a serious debilitating condition, which at present, is treated by symptomatic management of pain. Accidental injuries also lead to the development of OA. Cartilage degeneration is fundamental to the pathogenesis of OA. We propose to study the transcriptional control of MMP-1, a major enzyme involved in the degradation of articular cartilage. Expression of MMP-1 gene and the corresponding protein is markedly increased under osteoarthritic condition. Because cytokines like IL-1 and TNF-a increase expression of MMP-1 and biomechanical factors also influence its expression in osteoarthritic, unstable joints, the objectives are to understand what components of the promoter region of canine MMP-q gene are influenced by these factors. At present no data is available on canine MMP-1 gene regulation. This proposal is aimed towards understanding the regulation of canine MMP-1 gene expression in response to biomechanical stress and cytokines by isolating canine MMP-1 gene; identifying the regulatory elements in the promoter responsive to biomechanical stress and cytokines; and analyze MMP-1 expression in chondrocytes of articular cartilage from normal and osteoarthritic dogs with the intent to develop novel therapeutic drugs to combat this disease.

Progressive Retinal Atrophy (PRA)

Pending Grant No 2442. Investigation of Candidate Loci for Progressive Retinal Atrophy Principal Investigator. Simon Petersen-Jones, DVM, PhD; Michigan State University Sponsor(s) Belgian Sheepdog Club of America, Inc., Collie Health Foundation, Papillon Club of America, Tibetan Terrier Club of America

Abstract: Progressive retinal atrophy (PRA) is a common cause of blindness in purebred dogs. It is an inherited disease and there are several different forms resulting from defects in different genes. Most forms are due to a defect in a single gene and are inherited in an autosomal recessive manner. We have been investigating genes that could cause this disease and seeing if they are the cause of PRA in several breeds of dog. In one breed we have evidence that the PRA is caused by a defect at a genetic locus already known to be responsible for PRA in several other breeds. We have shown that this locus is not involved in PRA in the other breeds. We wish to investigate a further 50 genes to see if they are involved in PRA in any breeds. If we identify the gene defect that causes the disease in one or more breeds we will develop a DNA test for the defective gene. Such a test will allow breeders to eradicate PRA from their breed.

Reproduction

Active Grant No 2416: Systemic Inflammatory Response Syndrome in Canine Pyometra Principal Investigator: Boel Fransson; Washington State University

Sponsor(s): Bulldog Club of America Charitable Health Fund, Inc., Chow Chow Club, Inc., English Setter Association of America

Abstract: Early recognition and aggressive treatment of the Systemic Inflammatory Response Syndrome (SIRS) is imperative to avoid a fatal outcome. SIRS is not a disease in itself but is considered as the overwhelming response of the body to a cascade of inflammatory mediators, released after major trauma, pancreatitis, snake bite, heat stroke or bacterial infection of different organs. Pyometra, a common disease in intact female dogs, is considered associated with SIRS. Our cooperation with the Swedish University of Agricultural Science in Sweden, where female dogs are not routinely spayed, gives us an unique opportunity to receive large numbers of samples from dogs with pyometra. Blood samples from 50 dogs have been successfully transported to WSU. Detection of SIRS today is limited to recognition of clinical criteria, defined for SIRS, such as temperature, heart rate, and white blood cell count, but these criteria are not very specific. The proposed study will test if cytokine and acute phase protein levels in the blood are more specific detectors of SIRS than clinical criteria, which could lead to a more accurate and rapid detection of this syndrome. Rapid recognition would enable institution of aggressive treatment earlier in the disease, likely leading to reduced suffering and mortality of these dogs.

Respiratory Disorders

Active Grant No. 2401: Characterization of the Clinical Features of Idiopathic Pulmonary Fibrosis in the West Highland White Terrier

Principal Investigator: Brendan M. Corcoran, MVB, PhD; University of Edinburgh Sponsor(s): Westie Foundation of America, Inc.

Abstract: A chronic respiratory condition has been noted for some time in West Highland White Terriers, and typically affects middle to old-aged dogs resulting in reduced exercise ability, respiratory difficulty and coughing. Eventually the condition results in respiratory failure and the dogs have to be euthanized. The condition develops slowly and consequently the changes are often assumed by the pet owner to be attributable to normal aging. However, on chest auscultation distinct crackling sounds are heard which are indicative of significant lung abnormality. Many veterinarians are aware of this condition and often refer to it as Westie Lung Disease. However, we have recently described this disease in more detail and have evidence to suggest it is caused by scar tissue formation within the lung. The underlying cause is unknown and so we have termed the condition Idiopathic Pulmonary Fibrosis. Because we have limited information on this disease, the aim of our proposed study is to investigate the clinical aspects of this disease with a view to improving understanding of the disease and improve diagnostic accuracy. With that information we can move forward with studies to investigate the cause and explore novel therapeutic approaches to management of the disease.

Pending Grant No 2463. Evaluation of Pharyngeal Function in Dogs with Laryngeal Paralysis Prior to and After Unilateral Arytenoid Lateralization

Principal Investigator: MaryAnn Radlinsky; University of Georgia

Abstract: Laryngeal paralysis causes difficult breathing, which significantly limits activity and can be life threatening, in many large breeds of dogs. Surgical treatment maintains one side of the larynx, or voice box, in an open position. Coughing and gagging associated with eating and drinking and aspiration pneumonia may be present at the time of diagnosis or can develop after surgery. The complication rate associated with surgery in general has been reportedly low, but a recent publication quoted a 34 percent complication rate, with 28 percent complication rate following the most common surgical method. The purpose of this study is to determine if swallowing function is abnormal in dogs with laryngeal paralysis and to determine if surgery worsens this condition, thereby increasing the risk of aspiration pneumonia. Real-time radiographs of swallowing, endoscopy and evaluation of laryngeal muscle activity will be used to evaluate laryngeal and pharyngeal function in dogs affected with laryngeal paralysis prior to and after surgical correction. We will compare these results to young, healthy, large breed dogs and older large breed dogs that have no signs of respiratory or swallowing disease.

Rickettsial Disease

Active Grant No 2610: Molecular Approach to Determine the Spectrum of Rickettsial Disease in Dogs

Principal Investigator: Edward Breitschwerdt, DVM; North Carolina State University Abstract: Although Rickettsia rickettsii is a well-characterized cause of acute, severe and sometimes-fatal Rocky Mountain spotted fever (RMSF) in dogs and people, the role of other Rickettsia species as a cause of disease in dogs is yet to be established. During the late 1990s, we developed a method using polymerase chain reaction assay, or PCR, to detect the presence of DNA of rickettsial organisms in patient blood samples. This PCR method was developed under the currently accepted assumption that R. rickettsii represents the only clinically relevant Rickettsia species inducing disease in dogs in the United States. Therefore, species differentiation was not deemed necessary. Based upon recent clinical, serologic, and PCR data derived from dogs by our research group, this assumption appears to have been inaccurate. Initial work will focus on determining the gene sequence of isolates previously obtained in tissue culture from clinically ill dogs, sequencing rickettsial DNA amplicons detected by current diagnostic methods, and developing PCR techniques that will differentiate R. rickettsii from other known Rickettsia species. We believe that this approach will reveal previously unrecognized pathogenic rickettsiae. Results will impact diagnostic, therapeutic and preventative strategies affecting the health of dogs in North America and perhaps throughout the world.

Search and Rescue

Active Grant No. 2336: Medical Surveillance of Dogs Deployed to the World Trade Center and the Pentagon

Principal Investigator: Cynthia Otto, DVM, PhD; University of Pennsylvania Sponsor(s): American Kennel Club, AKC Companion Animal Recovery, Golden Retriever Foundation, Nestle Purina PetCare Company, PetSmart Charities, Veterinary Pet Insurance Company

Abstract: September 11, 2001, will live on in the memory and psyche of the American people. These events may also alter the lives of the hundreds of dogs and their handlers that served in a time of disaster. Federal (FEMA) Urban Search and Rescue Teams, police and private search dog handlers were involved in the search mission. The dogs were exposed to numerous hazardous materials. Although the acute medical problems were limited, it is impossible to predict the long-term effects of this disaster on the health and behavior of the dogs and the mental health of their handlers. In order to identify problems, we will perform intensive surveillance of the FEMA Team dogs and survey monitoring of the remainder of dogs. The intensive monitoring will include blood work (for evidence of infection, toxic injury and cancer) and chest radiographs over the next three years. Behavior and activity information will be collected at each time period for all dogs. The medical and behavior changes in the search dogs will be compared to controls to determine the lasting effects of this disaster and its response. Psychological effects on the FEMA dog handlers will also be monitored and interactions with the medical and behavioral effects of the dogs will be evaluated.

Active Grant No. 2337: Assessment of Injuries, Environmental Toxins and Anthrax Exposure in NYPD Search and Rescue and Bomb Detection Canines Du8ring World Trade Center Relief Efforts

Principal Investigator: Philip R. Fox, DVM; Animal Medical Center Sponsor(s): American Kennel Club, AKC Companion Animal Recovery, Nestle Purina PetCare Company

Abstract: Terrorist events of September 11 caused devastation of extraordinary scale and magnitude. There is little information in the scientific literature that speaks to the risks or medical consequences to search and rescue dogs exposed to environmental toxicities resulting from massive building collapse and incineration, and bioterror agents such as anthrax. We intend to identify and describe acute and chronic injuries, detect environmental-related toxicities, screen for zoonotic disease (anthrax), and assess future wellness and health risks of New York Police Department working patrol canines associated with the WTC disaster relief efforts. This knowledge will greatly facilitate current and future plans for emergency preparedness and management.

Skin Disease

Active Grant No 1815. Hair Coat Differences in Canine Breeds Recognized by the American Kennel Club

Principal Investigator Kelly Credille, DVM, PhD; Texas A&M University Sponsor(s): Anonymous

Abstract: We propose to define by picture and measurement the features that distinguish the different coat types of the most common breeds of dogs as recognized by the AKC. We do not know how the coat of a Whippet differs from a Beagle or what makes the coat of one dog better than another. In addition, defining normal in terms of number and diameter of hairs per unit area can be used to assess the severity of diseases with hair loss. This knowledge will allow veterinarians to make a definitive diagnosis of some diseases without the need for biopsy or laboratory tests. Finally, data obtained in this proposal can serve as a means of breed identification. A DNA bank of different canine breeds will also be assembled to provide a resource to further understanding of the canine genome and its diseases.

Active Grant No. 2272: Localization of the Gene for Sebaceous Adenitis in the Akita Dog by Homology Mapping

Principal Investigator: Kelly Credille, DVM, PhD; Texas A&M University Sponsor(s): Akita Club of America, Inc.

Abstract: Whenever I discuss with lay audiences the devastating effect genetic diseases can have on purebred dogs, I describe sebaceous adenitis (SA). A disease common in breeds like the Akita and Standard Poodle, severely affected dogs are largely hairless, covered in scale and smell terribly. There is no cure. Because the disease seldom shows itself until sexual maturity and because some affected dogs may never develop overt clinical signs, breeders cannot determine which dogs to breed. For these reasons, SA is a disease in need of a genetic test. However, designing such a test is challenging. Because of the difficulty distinguishing SA-affected from normal dogs, pedigree-dependent tests cannot be recommended. To accomplish the objectives of this study, we have devised an alternative protocol to associate a marker with SA. We have chosen the Akita breed to study because its history of bottlenecks and isolation make it ideal for this type of investigation. We believe this protocol will not only be successful but will result in a cost-effective and time-efficient alternative to more widely used methods in canine genetics research and will improve our ability to define those diseases in which distinguishing affected dogs can be ambiguous.

Active Grant No 2290 Mapping Canine X Chromosome Linked Alopecia Principal Investigator Gary S. Johnson, DVM, PhD; University of Missouri Sponsor(s) American Pomeranian Club, Inc., Anonymous, Pomeranian Philanthropists Abstract: Many young Pomeranians develop a luxurious puppy for first hair coat that fails to shed and is not replaced by an adult coat. As the puppy coat ages it breaks off and falls out and can result in a dog that is hairless over much of its body. This disease is sometimes called black skin disease, coat funk or woolly coat. It also occurs in Keeshonden and Alaskan Malamutes. Although females can have the disease, it is much more common in males. This suggests, but does not prove, that the mutation responsible for the disease is on the X chromosome. We propose to determine if a DNA marker from the canine X chromosome associates with the disease. If so, this marker could then be used to distinguish genetically normal puppies from

puppies that are likely to develop the disease. This marker could also identify female puppies that will not develop the disease but are likely to pass the disease on to the next generation.

Thyroid Disease

Pending Grant No. 2434 Recombinant Thyrotropin (TSH): Standard for the Next Generation of Canine TSH Immunoassays with Improved Sensitivity

Principal Investigator: Duncan Ferguson; University of Georgia

Sponsor(s): Airedale Terrier Club of America, Akita Club of America, Inc., American Belgian Malinois Club, American Boxer Charitable Foundation, American German Shepherd Dog Charitable Foundation, Borzoi Club of America, Clumber Spaniel Club of America, Collie Health Foundation, Dalmatian Club of America Foundation, Inc., English Setter Association of America, Golden Retriever Foundation, Italian Greyhound Club of America, Keeshond Club of America, Komondor Club of America, Miniature Pinscher Club of America, Inc., Norwegian Elkhound Association of America, Inc., Petit Basset Griffon Vendeen Club of America, Portuguese Water Dog Foundation, Rhodesian Ridgeback Club of the United States, Scottish Terrier Club of America Health Trust Fund

Abstract: Hypothyroidism, a failure of the thyroid gland, is the most common hormonal abnormality in dogs, causing a variety of medical problems in many breeds, including hair loss and skin infections. The measurement of serum levels of the pituitary hormone thyrotropin (TSH) has been used as a reliable and sensitive screening test for thyroid glandular insufficiency in human medicine for many years, but the "first generation" assays for canine TSH (cTSH) are missing as many as 1 out of 4 cases of hypothyroidism, resulting in no improvement in diagnostic sensitivity compared to total T4 measurement. Furthermore, the available assays have not been sensitive enough to distinguish low values of cTSH from those in the normal range. Towards the goal of improving current and future immunoassay sensitivity based upon a pure recombinant canine TSH (cTSH) hormone standard, our laboratory has succeeded in cloning and sequencing the two peptide subunits of canine TSH and have expressed them in small quantities. Using techniques recently developed in our parallel work on equine TSH, we plan to express and purify recombinant canine TSH in high quantities and validate its use as a pure immunoassay standard to facilitate its worldwide use.

Active Grant No. 2447. Genetic Determinants of Susceptibility to Hypothyroid Disease in Dogs Principal Investigator: George Happ, PhD, University of Alaska, Fairbanks Sponsor(s): Great Dane Club of America

Abstract: Canine hypothyroid disease is very similar to Hashimoto's disease in humans, which has been shown to be associated with human MHC genes. If we can show in hypothyroid digs a similar association with canine MHC genes, these could provide useful genetic markers for selective breeding to reduce disease incidence in purebred dogs. Hypothyroid disease is the most common endocrinopathy of dogs, and represents a significant veterinary problem. Definitive diagnosis is difficult since good clinical diagnostic tests are not available. Low levels of thyroid hormones characterize the disease, but these may result from other diseases and it appears that primary hypothyroid disease is characterized by the presence of autoantibodies to thyroglobulin. It is thought that only 50 percent of dogs with hypothyroid disease have these autoantibodies. We have recently developed an assay to measure thyroglobulin autoantibodies, which will allow

us to identify animals with primary disease. There is a clear genetic component to canine hypothyroid disease, and a number of breeds are thought to be more susceptible. The proposed study could lead to a better understanding of this condition and offer new approaches to its reduction in certain breeds.

Urinary Incontinence

Pending Grant No. 248 Use of an Artificial Urethral Sphincter in Dogs with Urethral Sphincter Mechanism Incompetence

Principal Investigator: Christopher A Adin, DVM, DACVS; University of Florida Abstract: Urinary incontinence (urine leakage) occurs in 13.6 percent to 20.1 percent of female dogs after elective spay procedures. Use of drugs to increase urethral tone are effective in many cases, though dogs that fail to respond to medical therapy require surgical intervention. Current surgical techniques have poor long-term success rates, with only 14 percent of dogs maintaining continence after one year in the most recent report. The purpose of this study is to investigate the use of a hydraulic urethral sphincter (HUS) in female dogs with urinary incontinence. Patient-controlled HUS have been used for many years in human beings and are highly effective in restoration of continence. Expense and complexity of design have prevented the patientcontrolled HUS from being applied in dogs. We have designed a simple, affordable HUS by combining an inflatable silicone cuff with a subcutaneous injection port. This silicone occluder is placed around the urethra via an abdominal incision, while the injection port is tunneled under the skin in the flank. Preliminary work has demonstrated that inflation of the occluder by injection of fluid into the subcutaneous port effectively increases resistance to urine leakage. Based on this preliminary data, we intend to evaluate the efficacy of the HUS in dogs with naturally occurring incontinence.

Vaccine Duration

Active Grant No 1604. Antibody Response and Duration of Immunity in Dogs Vaccinated with Attenuated Canine Distemper Virus, Canine Adenovirus Type 2 and Canine Parvovirus Vaccines from Commercial Sources

Principal Investigator. Saralyn Smith-Carr, DVM; Auburn University

Abstract: Annual revaccination of dogs with a combination canine distemper virus, canine adenovirus type 2 (infectious canine hepatitis), and canine parvovirus vaccine is routine although there is no scientific documentation to support the practice. There is evidence that current vaccines may provide long-term protection. If annual revaccination is not essential for the prevention of these infectious diseases, risk of vaccine-associated side effects could be avoided. The proposed study is to determine the minimal number of doses of vaccine necessary to immunize puppies and maintain protection in adulthood. Blood for assay of antibody levels will be collected from each of 180 healthy dogs (60 puppies, 12 weeks old or less and 120 adult dogs). Assays for antibody in blood serum will be performed on puppies prior to the first vaccination and two weeks following a second vaccination. Puppies with antibody levels sufficient to protect them against these viruses will require no additional vaccines in the initial

series. All dogs will be tested annually for three consecutive years. Additional vaccines will be administered as necessary to sustain immunity. Immunity will be measured by a standard assay and the data will be statistically analyzed. Antibody values that are essential for protection against these viruses have been reported. That information will be used to determine the criterion for acceptable antibody levels.

Mapping the Canine Genome

As breeders, we are challenged by patterns of inherited diseases in our dogs. Sometimes we can make sense out of these patterns; at other times disease seems to strike without warning in unexpected ways and at any age. We know we should not breed affected dogs. Anyone can unknowingly breed two carriers, however, and produce a dog with a disorder. The only way to eliminate serious disease is by developing tests that identify carriers. Once carriers are identified, we can avoid breeding these dogs together, thereby preserving the outstanding qualities of our breeds and breeding programs.

Developing a map of the canine genome is a major step forward in advancing the study of canine genetics and expanding our ability to identify genetic diseases. Healthier dogs and improving life for them and their owners is the mission of the AKC Canine Health Foundation. Over the last nine years, the Foundation has allocated more than \$1 million to canine genome map sequencing projects. In the last year, the National Institutes of Health have agreed to fund the canine genome-sequencing project at more than \$50 million. It is anticipated that the first stages of this map will be completed by the summer of 2004.

Active Grant No 2003: Anchoring the Canine Map Through BAC Mapping Principal Investigator: Elaine Ostrander, PhD; Fred Hutchinson Cancer Research Center Sponsor(s): Geraldine Rockefeller Dodge Foundation, Newfoundland Club of America Charitable Trust

Abstract: Genome maps are essential in identifying genes that cause inherited disease. They consist of a series of markers, positioned along each chromosome, which act as reference points for navigating different regions of the genome. Currently, the canine map is composed of about several hundred such markers, which provide "addresses" for over 90 percent of the genome. This early version of the map has proven useful for identifying the general location of several disease genes. But a much more highly refined map is necessary if we are to actually clone disease genes of interest (not just identify their location) and, subsequently, develop highly reproducible genetic tests. This proposal aims to characterize several hundred random clones, each containing a small portion of the canine genome, and then mapping them on the existing map relative to the markers and genes already in place. These clones, called BACs, will serve as "entry points" along the canine genome. Once a region of the genome is identified where a disease gene lies, the BAC clones will serve as starting points for investigators to genetically "walk" up and down the region and eventually clone the gene of interest. The work done as a result of this proposal is not breed specific, rather it will benefit all breeds of dogs equally.

Active Grant No. 2042. Identification and Mapping of Genes Expressed in the Canine Brain Principal Investigator: James R. Mickelson, PhD; University of Minnesota Abstract: The field of molecular genetics has produced remarkable medical advances in the last decade. For example, the underlying genetic basis for inherited diseases such as cystic fibrosis, muscular dystrophy, Huntington's disease, and certain forms of cancer have now been revealed. This research has produced sensitive tests for susceptibility to, and early diagnosis of, many types of diseases in humans, and the latest treatments based on this research have improved the quality of life and life span of numerous individuals. Molecular genetics is also contributing to an enhanced understanding of normal and disease processes in domestic and companion animals. Animal breeders have intuitively recognized for decades that certain traits in their animals may be inherited, and there is increasing awareness among breeders that genetics plays an important role in many other facets of animal health including disease susceptibility and resistance. In this proposal we hope to greatly increase the number of genes expressed in the canine brain and central nervous system whose DNA sequence and precise location on a canine chromosome is known. This work will help further efforts to identify those canine nervous system genes that greatly impact the health and other characteristics of all dogs by making the extensive human genome mapping information more directly transferable to knowledge of the canine genome.

Active Grant No. 2215. A BAC Map of the Canine Genome

Principal Investigator. Elaine Ostrander, PhD, Fred Hutchinson Cancer Research Center and Francis Galibert, PhD; CNRS, France

Sponsor(s): Geraldine Rockefeller Dodge Foundation

Abstract: Genome maps are essential for identifying genes that cause inherited disease. They consist of a series of markers, or molecular street signs, positioned along each chromosome, which act as reference points for navigating different regions of the genome. Currently, the canine map is composed of about several hundred such markers, which provide "addresses" for over 90 percent of the genome and as a result, the current map has proven useful for identifying the general location of several disease genes. But a much more highly refined map is necessary if we are to actually clone disease genes of interest (not just identify their location) and, subsequently, develop affordable and reproducible genetic tests. This proposal aims to characterize several hundred random clones, each containing a small portion of the canine genome, and then order them on the existing map, relative to the markers and genes already positioned. These clones, called BACs, will serve as "entry points" along the canine genome to begin a molecular "walk" towards a disease gene once its general chromosomal location is known. The work done as a result of this proposal is not breed specific; rather, it will equally benefit all breeds of dog.

Active Grant No. 2248 Generation of Canine EST Resources
Principal Investigator Richard W McCombie, PhD; Cold Spring Harbor Laboratory
Sponsor(s) Geraldine Rockefeller Dodge Foundation

Abstract: The identification of all of the genes, or major elements of heredity, within an organism is revolutionizing biomedical research. This holistic approach referred to as genomics will enable us to understand and eventually solve diseases with complex genetic factors underlying them. While genomic analysis of the dog is at an early stage, the impending completion of the mouse and human genome sequences will allow rapid acceleration of canine genomics. We are proposing to contribute to this process by building resources for gene

identification in the dog. This will include constructing libraries of genes from the dog and determining the DNA sequence of a large number of their component genes. These can then be compared to their counterparts from human and mouse.

Active Grant No. 2448: Identification and Mapping of Genes Expressed in the Canine Brain Principal Investigator: James R. Mickelson, PhD; University of Minnesota Abstract: Molecular and medical genetics is producing remarkable advances and contributing to an enhanced understanding of normal and disease processes in humans. We are now at the stage in canine genetics where we also hope to make rapid progress in identifying the molecular and genetic bases for traits that play a role in health, disease susceptibility, and disease resistance in dogs. Canine neurophysiology and behavior is ultimately governed by the expression of many thousands of genes in the brain and central nervous system. A number of devastating canine diseases are also due to either heritable or acquired alterations in the function of nervous system genes. In this study we will greatly increase the number of these important, interesting and useful genes whose precise location on a canine chromosome is known. This work will help future efforts to identify those genes that greatly impact the health, behavior, reproduction and other characteristics of all dogs by making the extensive human genome mapping information more directly transferable to knowledge of the canine genome.

America	n Kennel Club	Canine He	alth Found	ation Board Roster fo	r 2003			
Title	ErretName	MiddleInitia	LastName	Address 1	City	State	Zıp	Class
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	Wayne	E		PO Box 37941	Raleigh	NC	27627-7941	2007
Mr	Robert	L.	Kelly	PO Box 37941	Raleigh	NC	27627-7941	2005
Mr		<u>L.</u>	Arnold	PO Box 37941	Raleigh	NC	27627-7941	2007
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Mrs	Catherine	D	Adier	PO Box 37941	Raleigh	NC	27627-7941	2005
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Mrs	Pamela	Stephens	Butherus	PO Box 37941	Raleigh	NC	27627-7941	2006
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Mr	Alexander	F	Draper		Raleigh	NC	27627-7941	2003
Mr	Howard		Falberg	PO Box 37941		NC	27627-7941	2006
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Prof	Iris	Cornelia	Love	PO Box 37941	Raleigh	NC		2008
Dr	Asa		Mays	PO Box 37941	Raleigh	NC	27627-7941	2007
Mr	Thomas	L	Millner	PO Box 37941	Raleigh	NC	27627-7941	2008
Dr	William	R	Newman	PO Box 37941	Raleigh	NC	27627-7941	2006
Mr	Steve	Т		PO Box 37941	Raleigh	NC	27627-7941	2007
Dr	William	С	Truesdale	PO Box 37941	Raleigh	NC	27627-7941	2005
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Dr	C	Creston	Farrow	PO Box 37941	Raleigh	NC	27627-7941	
Mr	Thomas	J	Crowe	PO Box 37941	Raleigh	NC	27627-7941	
Dr	Robert	J	Hritzo	PO Box 37941	Raleigh	NC	27627-7941	
Staff								
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Ms	Erika	Α	Werne	PO Box 37941	Raleigh	NC	27627-7941	
Ms.	Diane		Vasey	PO Box 37941	Raleigh	NC	27627-7941	

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FINANCIAL STATEMENTS

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FINANCIAL STATEMENTS

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6300 Rockside Rd · Suite 100 · Cleveland, Ohio 44131 (216) 573-2330 · Fax (216) 573-0771 http://www.walthall.com

INDEPENDENT AUDITORS' REPORT

February 17, 2004

The Board of Directors of American Kennel Club Canine Health Foundation, Inc.

We have audited the accompanying statements of financial position of the American Kennel Club Canine Health Foundation, Inc (the "Foundation") as of December 31, 2003 and 2002 and the related statements of activities and changes in net assets, functional expenses and cash flows for the years then ended. These financial statements are the responsibility of the Foundation's management Our responsibility is to express an opinion on these financial statements based on our audits.

We conducted our audits in accordance with auditing standards generally accepted in the United States of America. Those standards require that we plan and perform the audit to obtain reasonable assurance about whether the financial statements are free of material misstatement. An audit includes examining, on a test basis, evidence supporting the amounts and disclosures in the financial statements. An audit also includes assessing the accounting principles used and significant estimates made by management, as well as evaluating the overall financial statement presentation. We believe that our audits provide a reasonable basis for our opinion.

In our opinion, the financial statements referred to above present fairly, in all material respects, the financial position of the Foundation as of December 31, 2003 and 2002 and the results of its activities and its cash flows for the years then ended, in conformity with accounting principles generally accepted in the United States of America.

Certified Public Accountants

Walthall, Drake & Wallace LLP

STATEMENTS OF FINANCIAL POSITION DECEMBER 31, 2003 AND 2002

	ASSETS		
		2003	2002
Cash and cash equivalents (Note 1)		\$1,079,337	\$ 966,590
Investments (Notes 1 and 2)		3,552,879	2,968,443
Dividends and interest receivable		10,212	20,203
Contributions receivable		357,653	458,776
Prepaid expenses		8,502	6,744
Furniture, fixtures and equipment, net (Notes 1	l and 3)	76,790	44,177
Charitable remainder annuity trust receivable (Note 1)	78,499	75,395
TOTAL ASSETS		<u>\$5,163,872</u>	<u>\$4,540,328</u>
	LIABILITIES		
Accounts payable		\$ 95,321	\$ 111,090
Grants payable (Notes 1 and 4)		2,446,818	2,095,744
Deferred contribution income			23,150
Amount due to broker			9,950
TOTAL LIABILITIES		2,542,139	2,239,934
	NET ASSETS		
Unrestricted (Note 1)		(695,841)	(773,059)
Temporarily restricted (Notes 1 and 6)		2,266,133	2,023,033
Permanently restricted (Note 1)		1,051,441	1,050,420
TOTAL NET ASSETS		2,621,733	2,300,394
TOTAL LIABILITIES AND NET ASSETS		<u>\$5,163,872</u>	<u>\$4,540,328</u>

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STATEMENTS OF ACTIVITIES AND CHANGES IN NET ASSETS FOR THE YEARS ENDED DECEMBER 31, 2003 AND 2002

		20	03	
	Unrestricted	Temporarily Restricted	Permanently Restricted	Total
REVENUES AND OTHER SUPPORT	<u> </u>			
Contributions	\$ 533,679	\$ 767,458	\$ 1,021	\$1,302,158
Contributions -		ŕ	,	
Interest and dividend income	97,237	3,104		100,341
Net unrealized and realized investment income (loss)	339,585			339,585
Corporate sponsored events and conferences (Note 8)	244,516			244,516
Administrative support - American Kennel Club, Inc. (Note 8)	220,000			220,000
Relocation support - American Kennel Club, Inc. (Note 8)	70,000			70,000
In-kind donation - American Kennel Club, Inc. (Note 8)	106,639			106,639
Royalty Income	8,993			8,993
Miscellaneous income	11,880			11,880
NET ASSETS RELEASED FROM RESTRICTIONS				
Satisfaction of program restrictions	527,462	(527,462)		
TOTAL REVENUES AND OTHER SUPPORT				
FUNCTIONAL EXPENSES				
Canine research and education	2,004,547			2,004,547
Fund raising	239,483		*-	239,483
Organizational development	134,969			134,969
General and administrative	503,774			503,774
TOTAL FUNCTIONAL EXPENSES	2,882,773			2,882,773
INCREASE (DECREASE) IN NET ASSETS	77,218	243,100	1,021	321,339
NET ASSETS - BEGINNING	(773,059)	2,023,033	1,050,420	2,300,394
NET ASSETS - ENDING	<u>\$ (695,841)</u>	\$2,266,133	<u>\$1,051,441</u>	\$2,621,733

2002					
	Temporarily	Permanently			
Unrestricted	Restricted	Restricted	Total		
\$ 294,278	\$1,033,012	\$ 1,017	\$1,328,307		
700,000			700,000		
116,594	2,982		119,576		
(349,348)			(349,348)		
255,078	20,500		275,578		
333,700			333,700		
13,163			13,163		
654,616	<u>(654,616)</u>				
2,018,081	401,878	1,017	2,420,976		
, ,					
2,053,720			2,053,720		
314,603			314,603		
196,402			196,402		
336,267			336,267		
<u> </u>					
2,900,992			2,900,992		
(882,911)	401,878	1,017	(480,016)		
109,852	1,621,155	1,049,403	2,780,410		
<u>\$ (773,059)</u>	<u>\$2,023,033</u>	<u>\$1,050,420</u>	<u>\$2,300,394</u>		

STATEMENTS OF FUNCTIONAL EXPENSES FOR THE YEARS ENDED DECEMBER 31, 2003 AND 2002

			2003		
	Canine Research	Fund	Organizational	General and	Total
	and Education	Raising	Development	Administrative	_Expenses
Grants (Note 1)	\$1,570,391	\$	\$	\$	\$1,570,391
Payroll and related expenses	226,485	73,854	68,679	67,951	436,969
Professional fees	45,943	12,218	12,094	103,194	173,449
Travel	9,167	11,188	1,383	7,904	29,642
Meetings	6,595	4,505	3,712	5,685	20,497
Conferences	64,474	69,936	12,168	, 	146,578
Printing, telephone, postage and office	34,717	14,421	13,603	5,732	68,473
Equipment rental and repairs	4,820	3,899	3,497	9,109	21,325
Rent and utilities	10,682	4,747	4,747	9,498	29,674
Marketing and advertising	1,659	19,972	3,000	18,974	43,605
Website design and expense	2,629	66	320	370	3,385
Membership expenses		2,376	2,171		4,547
New development		25	114		139
Promotional items purchased		8,083			8,083
Depreciation and amortization		4,001		15,345	19,346
Relocation expense				210,701	210,701
Retirement of assets				20,837	20,837
In-kınd donation (AKC) -					
office space and services	22,628	6,639	6,639	6,169	42,075
Miscellaneous	4,357	3,553	2,842	22,305	33,057
TOTAL	<u>\$2,004,547</u>	<u>\$239,483</u>	<u>\$134,969</u>	<u>\$503,774</u>	<u>\$2,882,773</u>

The accompanying notes are an integral part of the financial statements

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		2002		
Canine Research	Fund	Organizational	General and	Total
and Education	Raising	Development	Administrative	Expenses
\$1,624,926	\$	\$	\$	\$1,624,926
151,204	120,985	93,564	123,331	489,084
90,319	3,123	10,590	122,138	226,170
18,999	15,347	6,216	10,814	51,376
6,671	2,247	2,144	3,853	14,915
54,891	56,154	6,671		117,716
48,586	36,730	25,370	12,449	123,135
8,914	7,083	5,646	4,650	26,293
	<u></u>			
14,947	10,991	7,034	10,992	43,964
3,564	17,469	5,159	23,803	49,995
26,346	260	8,805	200	35,611
	15,616	15,616		31,232
	8,039	2,987		11,026
	5,014			5,014
	12,992	4,990	12,269	30,251
4,353	2,553	1,610	11,768	20,284
<u>\$2,053,720</u>	<u>\$314,603</u>	<u>\$196,402</u>	<u>\$336,267</u>	<u>\$2,900,992</u>

STATEMENTS OF CASH FLOWS FOR THE YEARS ENDED DECEMBER 31, 2003 AND 2002

	2003	2002
CASH FLOWS FROM OPERATING ACTIVITIES		
Increase (decrease) in net assets	\$ 321,339	\$ (480,016)
Adjustments to reconcile increase (decrease) in net assets		
to net cash provided by operating activities.	10.246	20.051
Depreciation and amortization	19,346	30,251
Net unrealized and realized investment (gains) losses	(339,585)	349,348
Non-cash contribution of securities	(12,317)	(10,209)
In-kind donation of furniture and equipment - AKC	(64,564)	
Retirement of furniture and equipment	20,837	
Changes in assets and liabilities:		
Dividends and interest receivable	9,991	4,363
Contributions receivable	101,123	(64,334)
Prepaid expenses	(1,758)	(980)
Accounts payable	(15,769)	79,076
Grants payable	351,074	307,063
Charitable remainder annuity trust receivable	(3,104)	(2,982)
Deferred contribution income	(23,150)	(18, 184)
Amount due to broker	(9,950)	<u>9,950</u>
Total adjustments	32,174	683,362
NET CASH PROVIDED BY OPERATING ACTIVITIES	353,513	203,346
CASH FLOWS FROM INVESTING ACTIVITIES		
Purchase of investments	(1,728,265)	(1,767,850)
Proceeds from sale of investments	1,495,731	1,672,615
Purchase of furniture and equipment	(8,232)	(10,686)
	(0.40.3(.6)	(105.001)
NET CASH USED IN INVESTING ACTIVITIES	(240,766)	(105,921)
INCREASE IN CASH AND CASH EQUIVALENTS	112,747	97,425
CASH AND CASH EQUIVALENTS - BEGINNING	966,590	869,165
CASH AND CASH EQUIVALENTS - ENDING	<u>\$ 1,079,337</u>	<u>\$ 966,590</u>
SUPPLEMENTAL DISCLOSURE OF CASH FLOW INFORMATION Cash paid during the year		
Interest	<u>\$</u>	<u>\$</u>
Income taxes	<u>s – </u>	<u>\$</u>

The accompanying notes are an integral part of the financial statements

NOTES TO FINANCIAL STATEMENTS DECEMBER 31, 2003 AND 2002

NOTE 1 - SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES

NATURE OF OPERATIONS

The American Kennel Club Canine Health Foundation, Inc. (the "Foundation"), established February 21, 1995, is a not-for-profit organization (exempt from Federal income taxes under Section 501(c)(3) of the Internal Revenue Code) formed for the purpose of furthering the advancement of knowledge of canine diseases and health care by clinical study, laboratory research and publication

BASIS OF ACCOUNTING

The financial statements of the Foundation have been prepared on the accrual basis of accounting

ESTIMATES

In preparing financial statements in conformity with generally accepted accounting principles, management makes estimates and assumptions that affect the reported amounts of assets and liabilities and disclosures of contingent assets and liabilities at the date of the financial statements, as well as the reported amounts of revenues and expenses during the reporting period. Actual results could differ from those estimates.

CONCENTRATION OF CREDIT RISK

The Foundation places its cash and cash equivalents with high-credit quality institutions. At times these balances may be in excess of the FDIC insurance limit. Cash and investments in money market funds and shares of registered investment companies are uninsured.

CASH AND CASH EQUIVALENTS

The Foundation considers demand deposits and all highly-liquid investments with a maturity of three months or less when purchased as cash and cash equivalents for the purpose of the Statements of Cash Flows

INVESTMENTS

Investments in mutual funds, commercial paper, marketable equity securities, and U.S. government obligations are stated at fair market value with both realized and unrealized gains and losses recognized in the Statements of Activities and Changes in Net Assets

CHARITABLE REMAINDER ANNUITY TRUST

The Foundation is a beneficiary under a charitable remainder annuity trust agreement under which the donor is entitled to annuity payments for the remainder of his life. Upon the donor's death, the assets in the trust will revert to the Foundation. The Foundation has reflected a receivable in the amount of \$78,499 and \$75,395 at December 31, 2003 and 2002, respectively, in its Statements of Financial Position, representing the present value of the future benefits to be received by the Foundation

NOTES TO FINANCIAL STATEMENTS DECEMBER 31, 2003 AND 2002

NOTE 1 - SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

CONTRIBUTIONS

The Foundation recognizes contributions received, including all unconditional promises to give, as revenues in the period received at their fair values. Conditional promises to give are recognized as revenues when the conditions on which they depend are substantially met. Temporarily restricted support is recorded as unrestricted support if the restriction is met in the same period as the support is received.

The Board of Directors has predicated funding for certain grants on receiving a stipulated amount of donor support. Pledges received on these grants are considered conditional pledges and are not included in revenue until the required donor support level has been obtained. As of December 31, 2003 and 2002, there was \$-0- of conditional pledges made on grants not meeting the required donor support level. Cash contributions received on these grants are shown on the Statements of Financial Position as Deferred Contribution Income.

CONTRIBUTIONS RECEIVABLE AND ALLOWANCE FOR DOUBTFUL ACCOUNTS

Contributions receivable reflected on the Statements of Financial Position are expected to be received within one year. Contributions receivable are stated at the amount management expects to collect from outstanding balances. Management provides for probable uncollectible amounts through a charge to operations and a credit to a valuation allowance based on its assessment of the current status of individual accounts. Balances that are still outstanding after management has used reasonable collection efforts are written off through a charge to the valuation allowance and a credit to contributions receivable. The Foundation considers contributions receivable to be fully collectible, accordingly, no allowance for doubtful accounts is required.

FURNITURE, FIXTURES AND EQUIPMENT

Purchased property and equipment are carried at cost Donated property and equipment are carried at the approximate fair value at the date of donation Depreciation is computed using primarily the straight-line method Depreciation charged to operations was \$19,346 and \$20,271 in 2003 and 2002, respectively.

MAILING LISTS

Mailing lists represented in-kind contributions to the Foundation and were recorded at their fair value at the date of contribution. Amortization was calculated on the straight-line method over three years. The lists were fully depreciated in 2002. Amortization charged to expense was \$-0- and \$9,980 in 2003 and 2002, respectively

GRANTS

Unconditional single or multi-year grants are considered incurred and charged to expense at the time of approval by the Board of Directors. Any grant cancellations approved by the Board of Directors are recognized at the time of approval.

The Board of Directors has predicated funding for certain grants on receiving a stipulated amount of donor support. These grants are considered incurred and charged to expense when the donor support level has been obtained. As of December 31, 2003 and 2002 there were \$-0- and \$373,317, respectively, of proposed grants not meeting the required donor support level.

NOTES TO FINANCIAL STATEMENTS DECEMBER 31, 2003 AND 2002

NOTE 1 - SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES (Continued)

NET ASSETS

Unrestricted net assets include contributions and investment income that will be used to fund canine research and educational programs designated by the Board of Directors Temporarily restricted net assets are those whose use has been limited by donors to a specific time period or purpose. Permanently restricted net assets have been restricted by donors to be maintained by the Foundation in perpetuity. Investment income from permanently restricted net assets is unrestricted.

ADVERTISING COSTS

The cost of advertising is expensed as incurred.

MEMBERSHIP EXPENSES

All costs related to acquiring new members are expensed as incurred.

FUNCTIONAL ALLOCATION OF EXPENSES

The costs of providing the various programs and activities have been summarized on a functional basis in the Statements of Activities and the Statements of Functional Expenses Accordingly, certain costs have been allocated among the programs and supporting services benefitted

NOTE 2 - INVESTMENTS

Investments in mutual funds, commercial paper, marketable equity securities and U S government obligations with readily determinable fair values are reported at their fair values in the Statements of Financial Position.

Investments are comprised of the following

	2003	2002
Mutual funds	\$1,959,960	\$ 991,867
Commercial paper	245,969	310,215
Marketable equity securities	245,131	332,554
U S government obligations	677,347	1,324,022
Certificates of deposit	424,472	
Other		9,785
	<u>\$3,552,879</u>	<u>\$2,968,443</u>

NOTES TO FINANCIAL STATEMENTS DECEMBER 31, 2003 AND 2002

NOTE 3 - FURNITURE, FIXTURES AND EQUIPMENT

Furniture, fixtures and equipment consisted of the following:

	_ 2003	2002
Furniture, fixtures and equipment	\$ 92,171	\$134,454
Less accumulated depreciation	(15,381)	(90,277)
Furniture, fixtures and equipment, net	\$ 76,790	\$ 44,177

Pursuant to the Foundation's relocation to the American Kennel Club's North Carolina operations center in June 2003, the Foundation retired fixtures and equipment totaling \$115,080 with an accumulated depreciation of \$94,243 In 2003, the American Kennel Club also donated \$64,564 worth of furniture, fixtures and facility improvements to the Foundation.

NOTE 4 - GRANTS PAYABLE

Grants payable are scheduled to be disbursed as follows.

2004	\$1,590,470
2005	<u>856,348</u>
	<u>\$2,446,818</u>

NOTE 5 - LINE OF CREDIT

The Foundation established a \$250,000 unsecured line of credit with a commercial bank in September 2003. Interest expense on the line of credit is the bank's prime rate plus three-quarter percent per annum (4.75% at December 31, 2003). There were no advances made on the line of credit.

NOTE 6 - TEMPORARILY RESTRICTED NET ASSETS

	2003	2002
Specific canine research	\$2,187,634	\$1,947,638
Annuity trust agreement (for future periods)	<u>78,499</u>	<u>75,395</u>
Total temporarily restricted net assets	<u>\$2,266,133</u>	\$2,023,033

Temporarily restricted net assets were released for research related to the study of specific diseases

NOTE 7 - LEASES

The Foundation leased office space under an operating lease agreement which was scheduled to expire in May 2004 In September 2003, a \$20,000 buyout of this lease was agreed to between CHF and the landlord. The buyout of the lease agreement is included in the relocation category on the financial statements. The Foundation also leases certain office equipment under an operating lease agreement which expires in September 2004.

The minimum future rent payable for this operating lease is \$4,743.

NOTES TO FINANCIAL STATEMENTS DECEMBER 31, 2003 AND 2002

NOTE 8 - RELATED PARTY TRANSACTIONS

consisting of a general contribution of \$800,000, administrative support of \$220,000, relocation support of \$70,000 and \$50,000 for the 2003 National Parent Club Canine Health Conference.	
During 2002, the general contribution of \$700,000, administrative support of) contributed funds totaling \$1,136,200 consisting of a of \$333,700 and \$102,500 for Search and Rescue research.
The Foundation relocated its offices to t	operation center in Raleigh, North Carolina in
June of 2003 In addition to the free use of office space	e, made improvements to the office space,
provided administrative support services and donated equi-	pment to the Foundation. The total estimated value of these
donated items was \$106,639	

) contributed to the Foundation funds totaling \$1,140,000

The Foundation's employees are covered under the Club's medical and pension plans as a related organization. The defined benefit pension plan, administered by the Club, is currently overfunded and therefore no pension expense is recorded in the accompanying Statements of Activities.

NOTE 9 - CONTINGENCIES

During 2003, the

The Foundation has received a claim for damages pursuant to a lawsuit initiated in June 2003. Management does not believe this matter will ultimately have a material adverse effect upon the financial condition of the Foundation and has not provided for this claim in the accompanying financial statements