#### 1 Risk and protective factors of feline tooth resorption in 8115 Finnish cats

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# 11 Abstract

12 Tooth resorption (TR) is the most common dental disease in cats. It causes painful lesions in the teeth, the only treatment being tooth extraction. The prevalence of the TR is high in clinical studies while 13 the estimate in the population sample is unknown. The aetiology of the disease remains unclear, but 14 associations to old age, breed, other oral and dental diseases, viral infections and certain 15 environmental factors have been suspected. We wanted to determine the prevalence, risk factors and 16 heredity of feline TR in a population sample of Finnish cats. We collected health and environmental 17 information of 8115 Finnish cats in 41 breeds through an online survey targeted for breeders and 18 owners. The prevalence of veterinary-diagnosed TR was 3.9% in the whole data and 15% in cats 19 20 diagnosed with oral or dental disease. Results indicated an increased risk by age and decreased risk 21 by constantly available food. Periodontitis and stomatitis were more common in the TR-affected cats. The interaction between gingivitis, dental calculus, and age suggests that the predisposition of young 22 cats to TR is associated with gingivitis, which could partially be prevented by proper dental hygiene. 23 The observed differences between breeds highlight the genetic contribution. 24

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# 26 Introduction

Tooth resorption (TR) is the most common dental disease in cats [1]. It is a painful disease 27 characterised by progressive dental destruction, which eventually results in loss of teeth. Dental 28 29 radiography is necessary to evaluate the overall situation [1]. Prevention of the disease is not possible since the aetiology is still unknown. The goal of treatment is to relieve the pain and discomfort caused 30 by these lesions [2]. The main cause of destruction are odontoclasts, which are multinuclear cells that 31 resorb mineralized tissue [3]. Odontoclasts are responsible for resorption of deciduous teeth in young 32 animals, but their abnormal activity in permanent teeth is the cause of TR [4]. The reason for this 33 34 process remains unclear, although many different theories have been proposed. Tooth resorption can

include plaque accumulation, inflammation of the adjacent tissue and alveolar bone ankylosis [5].
Because the pathogenesis of tooth resorption has been unclear, many different terms have been used
to describe the lesion, i.e. erosion, neck lesion, FORL (feline odontoclastic resorptive lesion).
Nowadays, most used term is TR.

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TR is a relatively new disease, which has become more common in the last decades [6]. Almost no 40 resorptions could be found in feline skulls from the 1930-1950's but starting from 1960's they were 41 increasing in number [7]. First publication about TR came out in 1955 [6]. The size and origin of the 42 study population have had a major impact on the reported prevalence of TR in previous studies 43 44 varying from 29-85% (Table 1). Most studies are based on a small clinical sample (n< 150) and no reports have been published yet in a large population sample. TR exists also in large cats, such as 45 leopards and lions, but rarely in humans and dogs [1]. The increasing number of tooth resorption 46 47 indicates a change in the environment of domestic cats [4]. For example, the factors preventing resorption in the outer surface of the teeth could have been compromised [1]. Only lesions that occur 48 in the crown or in the cemento-enamel junction can be detected clinically [1]. However, resorption 49 can occur at any surface of the teeth or in the root and can lead to massive destruction in the adjacent 50 51 tissue.

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Reference	Country	Target population	Study population	Method	Prevalence
			(n), average age		(%)
			(years)		
Coles 1990	Australia	General	64, NK	Clinical	52
[13]		anaesthesia			
van Wessum	Netherlands	Dental procedure	432, NK	Clinical	62
et al. 1992					
[7]					

**Table 1.** Prevalence of feline tooth resorption in previous studies. NK = not known

van Wessum	United	Dental procedure	78, NK	Clinical and	67
et al. 1992	States			radiographs	
[7]					
Lund et al.	United	General	145, 7.9	Clinical	48
1998 [15]	States	anaesthesia			
Ingham et al.	United	Clinically healthy	228, 4.9	Clinical and	29
2001 [16]	Kingdom	test animals		radiographs	
Pettersson &	Sweden	Sedation	96, 6.0	Clinical and	32
Mannerfelt				radiographs	
2003 [17]					
DeLaurier et	United	Clinically and	13, NK	Electron	85
al. 2009 [5]	States	radiographically		microscopy	
		normal teeth			
Our study	Finland	Finnish cat	8115, 5.3	Questionnaire	3.9 / 15
		population: all /			
		cats with oral or			
		dental disease			

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American Veterinary Dental College has developed a classification for tooth resorption based on their radiographic appearance. In type 1 tooth resorption there is a focal radiolucent area in the tooth that appears otherwise normal with a normal periodontal ligament space. In type 2 tooth resorption, there is decreased radiopacity, which indicates that the root is being replaced with alveolar bone. In type 3 tooth resorption, there are signs of both type 1 and type 2 resorptions [8,9]. The same cat can suffer from different types of resorptions [9]. The three resorption types have been suspected to have different aetiologies [10].

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Unlike the previous clinical studies, our aim was to investigate the prevalence of veterinary-diagnosed
TR, its risk factors and variation between breeds in a large population sample of Finnish cats collected
during 2012–2015 using an online feline health survey data [11].

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# 67 **Results**

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## 69 **Prevalence**

Our cat population consisted of 8115 cats in 41 breeds, of which 4290 (53%) were females. 2070 70 (26%) cats were diagnosed with oral or dental disease and 316 cats had veterinarian diagnosed TR. 71 The mean age of the cat population was 5.3 years - for cats without TR 5.1 and for cats with 72 veterinarian diagnosed TR 9.8 years. The prevalence of veterinary-diagnosed TR in our study was 73 74 3.9% (95% CI 3.5-4.3%, Table 1). The prevalence increased strongly by age, being 0.4% in 1 to <3 years old, 3.0% in 3 to <7 years old, 8.8% in 7 to <11 years old, and 11.6% in at least 11 years old 75 cats. TR was not reported in cats under one year of age. For comparison, in subgroup of cats that had 76 77 been diagnosed with oral or dental disease, the prevalence of TR was 15.2% (95% CI 13.5–16.8%, Table 1), increasing again by age 3.7%, 12.6%, 24.3% and 28.1%, respectively. 78

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We did not find any significant difference between the TR prevalence in purebred cats and non-80 pedigree house cats (3.8%; CI 3.4–4.3% and 4.2% and CI 3.3–5.3%, respectively, P = 0.551, Fisher). 81 82 However, in some breeds, the prevalence was much higher or lower than in the entire population or in-house cats – high in Siamese (9.9%), Abyssinian (9.3%), Oriental Shorthair (9.1%) and Cornish 83 Rex (8.9%), and much lower than average in Turkish Van (0.4%) and Birmans (1.3%). TR was not 84 85 reported in breeds of Burmilla, American Shorthair, Don Sphynx, Egyptian Mau, Kurilian Bobtail, Manx, Neva Masquerade and Seychellois. However, these breeds were present in low numbers in our 86 dataset. 87

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## 89 Factors associated with tooth resorption

Total of 15 factors were qualified into the final multivariable logistic regression modelling for finding 90 91 out the most significant factors associated to TR. They were age, breed, gender, availability of food, gingivitis, stomatitis, periodontitis, dental calculus, tooth fracture, abnormal number of teeth, cat flu, 92 musculoskeletal disease, digestive tract disease, respirator system disease and endocrinological 93 disease and tumours. These 15 factors were selected out of 51 (Supplementary Table S1) factors from 94 the feline health survey [11], which were known to be or could possibly be related to TR. 95 Qualification by basic association tests (Fishers' exact or Kruskal-Wallis, p>0.2) suppressed the 96 number of factors from 51 to 26 (Table 2, Supplementary Tables S2 and S3). Logistic regression 97 modelling with confounding factors age, gender and breed - for each of the 26 variables separately -98 99 favoured 15 factors (p <0.05) (Supplementary Tables S2 - S4) to be approved into the final multivariable logistic regression modelling. Majority of the cats - 92% of cases and 78 % of controls 100 - were neutered and thus, too small comparison groups of non-neutered cats prevented validation of 101 the effect of neutering in obtaining TR. 102

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104**Table 2.** Demographics and basic association test results (breed and disease categories excluded) of105cats with and without veterinarian diagnosed tooth resorption (N = 8115); number (n), percentage106(%) and association with tooth resorption (only P < 0.2 shown) in Finnish cats in feline health survey</td>10712/2012–2/2015. P-value: Fisher or Kruskal-Wallis test. CI = confidence interval. Bolded: selected108for further analysis in logistic regression model with confounding factors age, gender and breed. Same109cat may have had several of the oral/dental diseases or viral infections.

Variable	Diagnosed			Not dia	gnosed		
	n	%	95% CI	n	%	95% CI	P-value
Age	311			7631			<0.001
< 1 yr	0	0.0	0.0–1.2	1119	14.7	13.9–15.5	
1–2.99 yrs	8	2.6	1.3–5.0	1973	25.9	24.9 <del>–</del> 26.8	
3–6.99 yrs	76	24.4	20.0–29.5	2490	32.6	31.6 <del>-</del> 33.7	
7–10.99 yrs	115	37.0	31.8–42.5	1197	15.7	14.9–16.5	
≥11 yrs	112	36.0	30.9–41.5	852	11.2	10.5–11.9	
Gender	313			7742			0.184
Female	155	49.5	44.0-55.0	4135	53.4	52.3-54.5	
Male	158	50.5	45.0–56.0	3607	46.6	45.5 <del>-</del> 47.7	
Oral/dental disease							
Gingivitis	109	34.5	29.5-39.9	499	6.4	5.9–7.0	<0.001
Stomatitis	18	5.7	3.6-8.8	43	0.6	0.4–0.7	<0.001

Variable	Diagn	osed		Not dia	Not diagnosed			
	n	%	95% CI	n	%	95% CI	P-value	
Periodontitis	36	11.4	8.3–15.4	68	0.9	0.7–1.1	<0.001	
Dental calculus	187	59.2	53.7 <b>–</b> 64.5	1407	18.0	17.2 <b>–</b> 18.9	<0.001	
Tooth fracture	16	5.1	3.1 <del>-</del> 8.1	90	1.2	0.9–1.4	<0.001	
Abnormal number of teeth	12	3.8	2.2-6.5	63	0.8	0.6–1.0	<0.001	
Viral infections								
Cat flu	20	6.3	4.1–9.6	201	2.6	2.2-3.0	0.001	
Feline infectious peritonitis (FIP)	1	0.3	0.1–1.8	110	1.4	1.2–1.7	0.133	
Diet								
Cooked meat/fish	88	27.8	23.2-33.0	1812	23.2	22.3–24.2	0.067	
Availability of food	166	53.2	47.7–58.7	5287	68.9	67.9 <del>–</del> 70.0	<0.001	
Vaccinations								
Cat flu	201	70.3	64.7–75.3	4907	74.2	73.1–75.2	0.148	
Panleucopenia	239	83.6	78.8 <del>-</del> 87.4	5752	86.6	85.8 <del>-</del> 87.4	0.157	
Leukaemia	28	12.8	9.0–17.9	936	20.0	18.8–21.1	0.009	
Rabies	80	32.4	26.9–38.5	2649	46.0	44.7–47.3	<0.001	

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# 111 Multivariable logistic regression model

Seven variables remained in the final multivariable logistic regression model after model validation with backward selection and goodness of fit tests (Table 3). Based on the model, independent risk factors for veterinarian-diagnosed TR were breed, stomatitis and periodontitis whereas availability of food constantly had a significant protective effect.

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Breeds and breed groups with significantly higher risk for tooth resorption in comparison to house cats were oriental group (Siamese, Balinese, Oriental, Seychellois), British, Cornish Rex, European, Norwegian Forest Cat and Ragdoll. Turkish Van had significantly lower risk for TR. Those cats, which did not have food available all the time as well as cats, which had stomatitis or periodontitis were more prone to TR than other cats.

123	The second order interaction was found between age, gingivitis and dental calculus. The interaction
124	demonstrated that there was considerable variation in the risk to have TR in the groups indicated by
125	these interaction factors (Fig 1). Gingivitis was a significant risk factor for TR in age groups of $< 7$
126	years and $\geq 11$ years old cats, and the risk of having TR due to gingivitis was highest in the subgroup
127	of young cats (< 7 years) having gingivitis but not dental calculus (Table 3).
128	
129	Even though the P-value of the second order interaction was non-significant (0.0949) in this model,
130	it gathered the information of the three separate significant interactions explicitly and the model
131	achieved the best AIC and AUC values and McFadden index. The AIC was 2004. The AUC-value

- 132 for the ROC-curve was 0.870 (95% CI 0.850–0.890), which makes the predictive value of the model
- moderate [12]. The McFadden goodness of fit index was 0.255. The goodness of fit and predictive
- value tests for the final model were mainly good or moderate.
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**Table 3.** Multivariable logistic regression model of the risk factors for veterinarian diagnosed tooth resorption in Finnish cats in data collected during 12/2012-2/2015 in an internet survey. Only significant breeds are shown. B = logistic regression model coefficient, se = Standard Error, Wald = Wald test statistic, df = degrees of freedom, P = Wald's P-value, OR = odds ratio, CI = confidence interval for OR, ref = reference group.  $n_{cases}=308$  nand  $n_{controls}=7508$ . P-values of the first order interactions age x gingivitis, age x dental calculus and gingivitis x dental were 0.003, 0.0003 and 0.0002, respectively.

Variable	В	se	Wald	Р	OR	95% CI	df
Constant	-5.25	0.26	410.40	<0.001			1
Breed: House cat	ref						20
Siamese, Balinese, Oriental, Seychellois	0.72	0.25	8.25	0.004	2.05	1.26 - 3.36	1
British	0.65	0.32	4.23	0.040	1.92	1.03 - 3.58	1
Cornish Rex	1.14	0.25	20.39	<0.001	3.12	1.90 – 5.11	1
European	0.89	0.29	9.54	0.002	2.44	1.39 – 4.31	1
Norwegian Forest Cat	0.64	0.29	4.97	0.026	1.89	1.08 - 3.32	1
Ragdoll	0.66	0.32	4.27	0.039	1.93	1.03 - 3.61	1
Turkish Van	-2.28	1.02	4.98	0.026	0.10	0.01 - 0.76	1
Food available vs. no	-0.47	0.13	12.52	<0.001	0.63	0.49 – 0.81	1

Oral /dental diseases								
Stomatitis vs. no	0.98	0.36	7.28	0.007	2.65	1.31 – 5.39	1	
Periodontitis vs. no	1.29	0.26	25.32	< 0.001	3.63	2.20 - 6.01	1	
Interaction age x gingivitis x dental calculus			4.71	0.095				
Gingivitis vs. no	Age							
when:	yrs							
Dental calculus	< 7				2.55	1.34–4.83	1	
No dental calculus	< 7				15.11	7.49–30.47	1	
Dental calculus	7-<11				1.47	0.84–2.55	1	
No dental calculus	7-<11				2.03	0.71–5.81	1	
Dental calculus	≥11				2.27	1.23-4.20	1	
No dental calculus	≥11				3.76	1.57–9.01	1	

#### 143

**Figure 1.** Dental calculus and gingivitis by age groups in cats with and without tooth resorption. Demonstrative bar chart of subgroup formation in the interaction between age, dental calculus and gingivitis in the multivariable logistic regression model of the risk factors for veterinarian diagnosed tooth resorption (Finnish cats in data collected during 12/2012–2/2015 in an internet survey). Blue = percentage of cats with dental in the age group. Grey = percentage of cats with no dental calculus in the age group. Horizontal black

149 lines in blue and grey bars: proportion of gingivitis in cats with dental/without calculus under the line.

# 151 **Discussion**

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This is the first study of TR and its associated factors in a large population sample [11]. Tooth 153 resorption is a disease of high prevalence, unclear pathogenesis and various clinical presentations. 154 The prevalence of veterinarian diagnosed TR in this study (3.9%) was clearly lower than in studies 155 that have clinically evaluated the teeth, usually between 29-67%. The prevalence of TR in our 156 restricted group of cats diagnosed with oral or dental disease (15%) is closer to that found in clinical 157 studies. The mean age of the population was not particularly lower than in other studies and so the 158 age of the cat population does not explain the low prevalence. The selection of the research population 159 has a major impact on the results. If the target population consists of cats seeking for dental treatment 160 routinely or because of dental disease, the prevalence of TR is expectedly higher than in healthy cats. 161 The methods used in various studies to detect the TR also affect the prevalence. Most obvious lesions 162 163 are found in general examination without sedation, but dental calculus can distract the examination, nor can the gum line be explored. Since examination with dental explorer instrument under 164 anaesthesia only detects lesions in the cementoenamel junction or in the crown area, dental 165 166 radiographs increase the prevalence even higher when lesions in the root area are detected.

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In this study, the tooth resorption was diagnosed by the veterinarian, but those cats used as controls had not been examined for TR as in clinical studies. This explains partly the low prevalence of TR in this study. Commonly cat owners do not monitor their pets' teeth, nor have the cats regular health checks by veterinarian. For these reasons and due to the progressive nature and challenging diagnostics, TR may not have been noticed in the healthy cats. Open ended questions revealed that in most of the cats the tooth resorption had been found during routine dental calculus removal, dental radiography was used for diagnosis and most of the cats had teeth removed because of TR.

Our finding that prevalence of TR increases by age is in alignment with other studies 176 177 [5,6,13,14,15,16,17,18]. Together with age, risk to have TR was also associated with gingivitis and dental calculus, leading to different odds of getting TR based on differential exposure to these factors. 178 Gorrel's (2015) theory suggests that tooth resorption consists of at least two aetiologically different 179 diseases: inflammatory type 1 resorptions and idiopathic type 2 resorptions [19] Of particular interest 180 was that gingivitis was extremely high risk to TR in young cats that did not have dental calculus. The 181 182 pathophysiological logical path is that dental calculus causes gingivitis, which in turn predisposes to resorption. On the other hand, gingivitis can occur without dental calculus, and it is an interesting and 183 anticipated finding that if young cats have gingivitis (with or without dental calculus) they also have 184 185 an increased risk of resorption, which probably is inflammatory, i.e. type 1 resorption. Based on this, one might suspect that there are more inflammatory type 1 resorptions in young cats, and thus, regular 186 tooth cleaning and dental examination could help young cats to avoid development of this type of TR 187 188 in the early stage of life. However, the very high odds observed in young cats with no dental calculus may be an overestimation, as many of youngest healthy control cats without reported dental calculus 189 had not undergone clinical dental examination. The owner may not have noticed dental calculus or 190 gingivitis. 191

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In addition to gingivitis and dental calculus, stomatitis and periodontitis were risk factors for TR in our study. Previous studies have shown a connection between inflammatory type 1 resorptions and stomatitis [8], but not all studies have found the connection [20]. It has been suspected that chronic stomatitis caused by feline calici virus has an impact on development of TR [1]. Low number of cats with stomatitis hindered us to study this connection properly. Periodontitis has been related to TR also in previous studies [4] and it has been linked especially with type 1 tooth resorption [8], but also the inflammation caused by tooth resorption has been suspected to cause periodontitis [21].

The continuous availability of food had a protective effect on TR. If there is food available all the 201 202 time, the cat probably goes to eat more often. On the other hand, if the food is available only on certain times, the cat might eat faster and more eagerly, which can also affect the teeth. The effect of 203 the availability of food has not been, to authors' knowledge, studied before. DuPont & DeBowes 204 (2002) suspected dry food to cause mechanical trauma leading to type 2 tooth resorption and soft 205 food, causing periodontitis, being a risk factor for type 1 resorption [8]. Scarlett et al. (1999) did not 206 207 find a difference in prevalence between cats eating dry or soft food [4]. In our questionnaire we did not specifically ask if the cats ate dry or soft food, but we did not find association between TR and 208 eating cooked meat/fish. 209

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Our finding of gender not being in association with TR was in line with previous studies [4, 13]. In 211 previous studies, indoor cats have been suspected to have higher risk of TR than outdoor cats [4], but 212 213 some studies have not found a difference [16]. We did not find difference in outdoor habits of the cats with or without TR and could not reliably evaluate the effect of neutering and vaccinations due to 214 uneven groups in neutering status and missing values in vaccinations. Considering other diseases, 215 feline infectious peritonitis (FIP), cat flu (including herpes and calici virus), leukaemia virus (FeLV), 216 217 immunodecifiency virus (FIV) and feline panleukopenia virus were not associated with TR in our 218 study.

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Varying results have been published considering the prevalence of TR in different breeds. Our results
revealed that TR is highly associated with breed – being purebred was not a risk factor as itself, but
the associations concerned certain breeds or breed groups. We found oriental group (Siamese,
Balinese, Oriental, Seychellois), British, Cornish Rex, European, Norwegian Forest Cat and Ragdoll
in higher risk for TR. Interestingly, the only breed having less TR than our comparison group of house
cats was Turkish van. Siamese, Cornish Rex and European have been found to be predisposed to TR

also before [7,11,13]. The breed predisposition might indicate a genetic component in the aetiology 226 227 of the disease. Some breeds might be genetically predisposed to dental and oral diseases in general. Exposition to several dental and oral diseases at the same time in cats have been observed in our 228 previous study [11], in which, for example Cornish Rex and the oriental group (Siamese, Balinese, 229 Oriental, Seychellois) were predisposed to periodontitis and stomatitis at the same time. However, it 230 seems that not all breeds that were associated with TR have the predisposition to other dental and oral 231 232 diseases. In our study, European was associated with TR whereas the previous study did not find European to be predisposed to other dental diseases [11]. The causalities between dental and oral 233 diseases are unclear, as they can occur on their own or with other oral diseases. Breeder's influence 234 235 on the prevalence of tooth resorption, stomatitis and periodontitis in the breed has also been suspected [11]. This is an interesting speculation, since this can indicate heritability or similarity in 236 environmental factors and infections. Further analysis of the genetic background of tooth resorption 237 238 would require a clinical trial where the type of resorption and dental health of the controls could be determined via radiography. 239

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Because the data were collected in a cross-sectional manner, only the effect of permanent risk factors such as breed, age and gender can be considered causal, since they have been permanently present before the disease. However, other factor found significant are potential risk or protective factors that should be verified with clinical trials or observational follow-up studies.

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# 246 Conclusions

We have performed the first large scale population study of TR and identify several predisposing factors. Old age, stomatitis and periodontitis were risk factors for TR. Together with age, risk to have TR was associated with gingivitis and dental calculus, suggesting that young cats that are susceptible to tooth resorption associated with gingivitis could be helped with proper dental hygiene. In addition, keeping food available constantly was a potential protecting factor. Finally, certain breeds appeared
more predisposed to TR, suggesting a genetic contribution to the aetiology of the disease.

253

# 254 Materials and Methods

The material for this study was part of a cross-sectional online feline health survey targeted to all 255 Finnish cat breeders and owners. The data originated from survey responses between December 2012 256 to February 2015 and included 8115 cat owners' responses from the feline health survey. The cats 257 258 belonged to 40 different breeds and non-pedigree house cats. We used sample size requirements for breeds as described by Vapalahti et al [11]. The sample size requirements were met in purebred cats 259 by 13 single breeds and 6 breed groups that consisted of 18 different breeds. The questionnaire 260 included information about the cats themselves, their environment, diseases and behaviour. The 261 diseases were divided into various categories and each category included information whether the 262 263 diagnosis was made by a veterinarian or by the owner. The content, the questionnaire and data collection methods in detail and some results of the survey have been published previously 264 [11,21,22,23]. Our data consisted of 1) basic information of the cat: breed, registration number, day 265 266 of birth, possible day of death, gender, neutering, 2) environmental factors: vaccinations, outdoor habits, diet, home environment, and 3) disease categories and specific diagnoses, of which only oral 267 and dental diseases, autoimmune diseases and viral infections were considered. The differential 268 269 diagnoses in dental and oral diseases were malocclusion, gingivitis, stomatitis, periodontitis, tooth resorption (in the survey feline odontoclastic resorptive lesion FORL), dental calculus, tooth fracture 270 and abnormal number of teeth. 271

The age of the cat was determined by the date of birth and the date of response. Age was categorized into age groups: <1 year, 1 to <3 years, 3 to <7 years, 7 to <11 years, and at least 11 years. For multivariable analysis the following age groups were used: <7 years, 7 to < 11 years, and at least 11 years. The gender of the cats was coded as 'male (1) and 'female' (2) and neutering as 'yes' (1) and

'no' (0). The association of TR with other conditions was studied both at the level of disease 276 277 categories and at specific diagnosis level. As an exception to other disease categories, in dental and oral disease category only the veterinarian's diagnoses were included in the study. In other disease 278 categories, the initial options 'veterinarian's diagnosis' or 'own diagnosis' were summed up to option 279 'yes' (1) if either option was selected. 'Not known' responses were coded as missing. Finally, 280 reencoding included two options: 'ves' (1) and 'no' (0). In the analysis of specific diagnoses, coding 281 282 'yes (1)' or 'no (0)' was used. The environmental questions, such as diet and vaccinations, were coded 283 similarly.

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285 Those cats that had been responded to have veterinarian diagnosed tooth resorption in the health survey, were determined as TR-cats. When evaluating breed associations, we used house cat as the 286 reference group. Even though logistic regression analysis is not the primary method to be used in data 287 288 based on cross-sectional study, it gives reliable results when the prevalence of the issue (here TR) is less than 5% [24] (3.9% in our study). Demographic and environmental factors of cats were examined 289 290 and tabulated. The prevalence of veterinarian diagnosed tooth resorption in different breeds and age groups was calculated with cross tabulation. Cross tabulation was used also to evaluate the 291 associations of TR with gender, other diseases and environmental factors. The 95% confidence 292 293 intervals for prevalence was calculated with Epitools [25] using the Wilson method [26]. Statistical significance of associations in cross tabulation were evaluated with Fisher's exact test for two-294 categorical variables, Chi-square test for multicategorical and Kruskal-Wallis test for ordinal 295 variables. After cross tabulations, variables at level P < 0.2 in basic tests were further analysed 296 individually in multivariable logistic regression with confounding factors. According to literature, 297 age, gender and breed were considered as confounding factors [27]. 298

Multivariable logistic regression modelling was performed to find the best model with the most 300 301 important risk factors. Interactions until the second order and multicollinearity between variables were tested. The model selection was performed by backward selection and goodness of fit statistics. 302 P-value <0.05 of Wald chi-square was set to cut off value for significance. Goodness of fit of the 303 model was evaluated with McFadden index and Akaike information criterion (AIC) and the predictive 304 value by the area under the curve (AUC) of the receiver operating characteristic (ROC) curve. 305 Multicollinearity between variables was estimated by Phi coefficient - the limit value for strong 306 correlation was set at 0.5 [28]. Microsoft Office Excel 2010 was used for data editing and IBM SPSS 307 Statistics, Version 22-24 for statistical analysis and SAS version 9.4, SAS Institute Inc., Cary, NC in 308 final logistic regression modelling. 309

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# 311 Ethical statement

The data in this study was collected 2012-2015 using an online feline health survey as published by Vapalahti et al.<sup>11</sup>. The data was collected before the onset of the GDPR regulation according to the Finnish legislation <u>https://www.finlex.fi/fi/laki/ajantasa/1999/19990523</u>. This survey study focused on the investigation of cats and not human participants or the cat owners, and therefore a specific ethical approval was not needed. As for the study participants (cat owners), we collected only names and addresses. Owners were informed that the participation is voluntary, confidential, and that the data is used only for scientific purposes. We received informed consents from all participants.

319

# 320 Additional Information

## 321 Acknowledgements

We thank the cat owners who participated in the original health survey.

#### 323

## 324 **Competing Interests**

- 325 The authors have declared that no competing interests exist.
- 326

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- 384 Author contributions
- 385 H.N., A-M.V. K.V. and H.L conceptualized and designed the experiment. H.N. made the preliminary
- analysis and K.V. performed the multivariable analyses. K.V. and H.N. drafted the manuscript, which
- 387 was edited and contributed by H.L and A-M.V. All authors approved the final version of the
- 388 manuscript.
- 389

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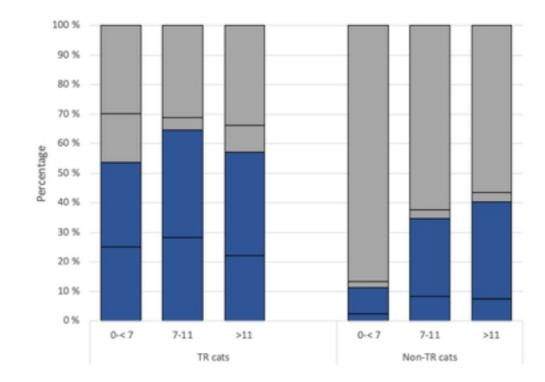
# **Supporting information**

- 391 Data available: Supplementary Tables S1-S4 and S5
- 392

# 393 Figure Legends

- 394
- **Figure 1.** Dental calculus and gingivitis by age groups in cats with and without tooth resorption.
- 396 Demonstrative bar chart of subgroup formation in the interaction between age, dental calculus and

397 gingivitis in the multivariable logistic regression model of the risk factors for veterinarian diagnosed 398 tooth resorption (Finnish cats in data collected during 12/2012–2/2015 in an internet survey). Blue = 399 percentage of cats with dental in the age group. Grey = percentage of cats with no dental calculus in 400 the age group. Horizontal black lines in blue and grey bars: proportion of gingivitis in cats with 401 dental/without calculus under the line.



# Figure 1