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### Otitis media and interna with or without polyps in cats

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1 **OTITIS MEDIA AND INTERNA WITH OR WITHOUT POLYPS IN CATS:**  
2 **ASSOCIATION BETWEEN MENINGEAL ENHANCEMENT ON POST-**  
3 **CONTRAST MRI, CSF ABNORMALITIES AND CLINICIAN TREATMENT**  
4 **CHOICE AND OUTCOME**

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47           Objectives: To evaluate the association between meningeal enhancement (MgE)  
48 and cerebrospinal fluid (CSF) analysis results, their individual association with  
49 bacteriology results from affected ear samples, and whether these test results influenced  
50 clinicians' therapeutic choice in cats with otitis media and interna (OMI).

51           Methods: Multicentre retrospective study over an eight-year-period. Cats  
52 diagnosed with OMI, with or without a nasopharyngeal polyp, leading to peripheral  
53 vestibular signs were included. Only cats for which MRI with post-contrast T1 weighted  
54 sequences and CSF analyses were available were included in the study. Cats with intra-  
55 axial MRI lesions or empyema were excluded.

56           Results: Fifty-eight cats met the inclusion criteria. MgE was reported in 26/58  
57 cases of which 9 had an abnormal CSF result (increased total nucleated cell count (TNCC)  
58 or total protein); 32/58 cases had no MgE of which 10 showed abnormal CSF results.  
59 There was no association between bacteriology results (external ear canal or bulla) and  
60 MgE or abnormal CSF results. CSF abnormalities were more commonly detected in acute  
61 cases (16/37) compared to chronic cases (3/21), the difference being statistically  
62 significant (Fischer test,  $P = 0.04$ ). Prednisolone was prescribed in 10/16 cases with  
63 increased TNCC. Among the 42 cases with normal TNCC, 15/42 received prednisolone,  
64 and 13/42 received non-steroidal anti-inflammatory drugs. Various antimicrobial drugs  
65 were prescribed in 53/58 cats. Antimicrobial therapy duration was similar regardless of

66 positive bacterial culture (5.58 vs 4.22 weeks), increased TNCC (6.13 vs 4.72 weeks) or  
67 MgE (5.33 vs 4.90 weeks).

68 Conclusion and relevance: No association was found between CSF and MgE  
69 results. Furthermore, no association was found between MgE, CSF or bacteriology  
70 findings, respectively. In addition, abnormal CSF results might lead the clinician to treat  
71 with corticosteroids but did not have any impact on antimicrobial therapy length. CSF  
72 abnormalities are seen significantly less frequently in chronic cases. Outcome tended to  
73 be poorer when MgE is detected on MRI.

74

75 Keywords: Otitis media and interna; Peripheral vestibular signs; MRI, Meningeal  
76 enhancement; Cerebrospinal fluid

77 **Introduction**

78 Otitis media and interna (OMI) has been reported to be the cause of cause  
79 vestibular signs in 43% to 63% of cats with peripheral vestibular signs (PVS) (1,2). OMI  
80 in cats is most commonly associated with inflammation caused by upper respiratory  
81 infection that has extended through the auditory tube or a nasopharyngeal polyp (1,3,4).  
82 It occurs less frequently as a consequence of an otitis externa or neoplasia. Bacterial  
83 isolates from OMI include *Staphylococcus* species, *Streptococcus* species, *Pasteurella*  
84 *multocida*, *Escherichia coli*, *Enterococcus* species and less frequently *Mycoplasma*  
85 species, *Corynebacterium* species and other bacterial species (1,4–8). Apart from  
86 bacteria, cats can have other infectious agents being involved, especially fungi  
87 (*Malessezia*) (8). Occasionally a nasopharyngeal polyp, a non-neoplastic, inflammatory  
88 growth that arises from the middle ear or the auditory tube, can be responsible for OMI  
89 (1,2,9,10). Diagnosis of otitis media and bacterial infection in the bulla can be reached  
90 via cytological and/or bacterial culture of material retrieved via myringotomy or surgical  
91 procedure such as bulla osteotomy (1,8,9). Cats with PVS are clinically recognized with  
92 the presence of at least one of the following clinical signs: ipsilateral head tilt, jerk  
93 nystagmus, tight circling, positional strabismus and/or vestibular ataxia and the absence  
94 of any neurologic signs suggestive of intracranial disease. (11–13). PVS reflect the  
95 involvement of the inner ear, while presence of facial nerve deficit and/or Horner  
96 syndrome indicates the involvement of the middle ear (11–13).

97 MRI is a sensitive method to diagnose OMI, particularly for inner ear visualization  
98 (1,7,14). The fluid composition of endo- and perilymph allows a good visualization of  
99 this inner ear part in a fluid sensitive sequence such as T2-weighted (T2W) images or in  
100 fluid-attenuated inversion recovery (FLAIR) (14,15). A marked hyperintensity compared  
101 to adjacent structures is present on T2W images while a suppression is visible in FLAIR  
102 (14,15). Post-contrast T1-weighted images may show abnormality consistent with  
103 inflammatory changes in the inner ear. (1) Typical changes raising the suspicion of OMI  
104 include isointense material in the bulla on T1W images and hyperintense on T2W images.  
105 On post-contrast T1W images, a peripheral enhancement along the inner surface of the  
106 tympanic bulla can be observed (7,16). A laminated appearance of the mucosa of the  
107 tympanic bulla on T2W images has also been described (17). A reduced signal intensity  
108 on T2W or an increase signal intensity on FLAIR images from the intralabyrinthine fluid  
109 are MRI finding suggestive of otitis interna (14,16,18). Due to anatomical proximity,  
110 intracranial extension of OMI can lead to a meningeal enhancement (MgE) on MRI after  
111 intravenous administration of paramagnetic contrast medium (19). Anatomically,  
112 perilymph and cerebrospinal fluid (CSF) are connected via the cochlear aqueduct (20).  
113 Therefore, CSF analysis is another important diagnostic tool for cats with OMI and is  
114 reported to be more sensitive than MRI to identify intracranial inflammatory processes  
115 (7,15,21). Thus, an abnormal MRI and/or CSF analysis can provide useful clinical  
116 information on the presence of a concurrent meningitis and may influence the therapy.

117           The relationship between MgE and abnormal CSF in cats with OMI has not yet  
118 been investigated, neither have their association with bacteriology results and therapeutic  
119 management. The aims of this study were to describe the association between MgE and  
120 CSF analysis results, their individual association with bacteriology results from affected  
121 ear samples, and the influence of the above with therapeutic choice in cats with OMI. We  
122 hypothesize that (a) MgE is associated with CSF abnormalities, (b) positive bacteriology  
123 is more common if MgE and/or CSF abnormalities are present and (c) positive  
124 bacteriology is associated with the choice and length of antimicrobial and/or anti-  
125 inflammatory therapy.



126 **Materials and Methods**

127 **Selection criteria**

128 Data were retrospectively collected from six different referral centres across  
129 Europe (Vetsuisse-Faculty, University of Bern; University of Veterinary Medicine  
130 Hannover, Foundation; Royal Veterinary College London (RVC); School of Veterinary  
131 Medicine, University of Glasgow; Queen's Veterinary School Hospital, University of  
132 Cambridge, and Royal (Dick) School of Veterinary Studies, University of Edinburgh)  
133 over an eight-year-period (January 2012 to December 2020). Only client-owned cats with  
134 peripheral vestibular signs that underwent MRI (with post-contrast images) and CSF  
135 analysis as part of the diagnostic work up were selected for this study. Inclusion criteria  
136 were: (1) clinical signs consistent with peripheral vestibular lesion localisation, (2) a  
137 diagnosis of otitis media and interna with or without the presence of a nasopharyngeal  
138 polyp based on MRI findings, (3) absence of intra-axial abnormality or imaging findings  
139 consistent with empyema on MRI.

140 Retrospective information collected from the medical records included  
141 signalment, history, therapy prior and after referral, side of PVS, MRI and CSF findings,  
142 bacteriology results from affected ear, and outcome.

143 **MRI**

144           The MRI images of the skull were obtained under general anesthesia, using  
145 anesthetic protocols at the discretion of the anesthesiologist in charge. High-field MRI  
146 were used at all institutions (except for one cat) and varied between centres: a Philips  
147 Panorama HFO 1.0 T (Philips Medical Systems Nederland B.V., Best, Netherlands) for  
148 Bern, a Philips Achieva 3.0 T (Philips Medical Systems, Best, The Netherlands) for  
149 Hannover, 1.5T (Intera; Philips Medical Systems, Amsterdam, Netherlands) for RVC,  
150 1.5T (Magnetom Essenza Siemens) for Glasgow, 0.27T (Esaote VetMR Grande, Genova,  
151 Italy) or 1.5 T (Phillips Achieva, Phillips Healthcare, Best, Netherlands) for Cambridge  
152 and 1.5 T (Intera; Philips Medical Systems, Amsterdam, Netherlands) for Edinburgh.  
153 Although MRI protocols varied between centres, in all cats, at least T1-weighted T1W,  
154 T2W and T1W post-contrast images were available (IV of 0.2 mmol/kg gadoteric acid  
155 [Dotarem; Guebert Laboratories] for Bern and Hannover, IV of 0.1 mmol/kg gadoterate  
156 meglumine [Dotarem; Guerbet, Milton Keynes, UK] for RVC, [Gadovist, Bayer] for  
157 Glasgow & Cambridge and 0.1 mmol/kg gadopentate dimeglumine [Magnevist, Bayer]  
158 for Edinburgh). All MR images had been evaluated by a board-certified veterinary  
159 radiologist or a board-certified veterinary neurologist. Information about OMI with or  
160 without nasopharyngeal polyp and MgE were collected directly from MRI reports.

161

162 **CSF analysis**

163 Abnormal CSF was defined as a total nucleated cell count (TNCC)  $\geq 5$  leukocytes  
164 per  $\mu\text{L}$  and/or increased total protein  $> 0.3$  g/L for cisterna magna samples and  $> 0.45$  g/L  
165 for lumbar samples. Albuminocytologic dissociation was defined as an increase in total  
166 protein without an increased TNCC. Neutrophilic (respectively monocytic) pleocytosis is  
167 identified if there is  $> 70\%$  of neutrophils (respectively monocytes) in a CSF with  
168 abnormal TNCC. (22)

169

170 **Medical treatment**

171 Antimicrobials were categorized as first or second line of treatment. First line  
172 included amoxiclav, cephalosporin, metronidazole, clindamycin and doxycycline.  
173 Second line contained marbofloxacin, enrofloxacin, cefovecine, pradofloxacin and  
174 cefixime. (23)

175 Anti-inflammatory drugs were categorized as steroidal (eg, prednisolone,  
176 dexamethasone) or non-steroidal (eg, meloxicam, robenacoxib).

177

178 **Statistical analysis**

179           The presence or absence of MgE on MRI, the CSF results and bacteriology culture  
180 results were compared using a  $\chi^2$  test. The choice of therapy based on MgE, CSF and  
181 bacteriology results were compared using a  $\chi^2$  test or a Fischer's exact test if a group  
182 contained less than 5 cats. Duration of therapy was compared using a Student's t-test. Test  
183 values were performed two-sided and a value of  $P \leq 0.05$  was considered statistically  
184 significant. All analyses were performed using R version 3.6.3.

185 **Results**

186 A total of fifty-eight cats met the inclusion criteria. Domestic shorthair cats were  
187 the most common breed (32/58 cats). Other breeds were Maine Coon (9), Siamese (4),  
188 British Short Hair (2), Burmese (2), Russian Blue (2), Bengal (1), Egyptian Mau (1),  
189 Norwegian (1), Ocicat (1), Persian (1), Ragdoll (1), and Snowshoe (1). There were 27  
190 females (four intact, 23 spayed) and 31 males (four intact, 27 neutered; sex ratio  
191 male/female = 1.15). The mean age of the cats was 6.9 years (median = 7.3 years, range:  
192 3.7 months to 14.7 years). Half of the cases had left vestibular signs at time of  
193 presentation. Clinical signs were acute ( $\leq 14$  days) in 37 cats and chronic ( $> 14$  days) in  
194 21 cats. Medical treatment before presentation was given in 32 cats. Nine cases received  
195 antimicrobial therapy alone, eight had an antimicrobial therapy with non-steroidal anti-  
196 inflammatory drugs, nine had an antimicrobial therapy with corticosteroids, and two had  
197 an antimicrobial therapy, non-steroidal anti-inflammatory drugs, and corticosteroids. Five  
198 cats received only corticosteroids and one only a non-steroidal anti-inflammatory drug.  
199 All clinical information from cases are available in supplementary material.

200 Forty-five cats (78%) were diagnosed with OMI alone (without a nasopharyngeal  
201 polyp) (Figure 1 (a), 2 (a) and (c)): three based on histology of material obtained from  
202 bulla osteotomy, three based on findings during bulla osteotomy, 26 based on otoscopy  
203 and cytology results obtained through myringotomy and 13 were suspected on MRI only.  
204 The remaining cats (13/58, 22%) were diagnosed with OMI secondary to a polyp (Figure

205 3 (a) and 4 (a): six based on histology of material obtained from bulla osteotomy, two  
206 based on findings during bulla osteotomy, two based on otoscopy and/or cytology results  
207 obtained through myringotomy and two were suspected on MRI only.

208

209 Meningeal contrast enhancement was present in 26/58 cases (45%) (Figure 2 (b)  
210 and 4 (b)) while 32/58 cases (55%) did not show MgE (Figure 1 (b) and 3 (b)). CSF  
211 analysis was abnormal in 19/58 (33%) cats. Eleven of them with only increased TNCC,  
212 five with both increased TNCC and total protein and three with albuminocytologic  
213 dissociation. Increased TNCC ranged between five and 1205 leukocytes /  $\mu$ L (median =  
214 12) and increased total protein in CSF ranged between 0.34 and 0.77 g/L (median = 0.56).  
215 Neutrophilic pleocytosis was seen in nine cases, monocytic pleocytosis was seen in one  
216 case and mixed cell pleocytosis in six cases.

217 Nine of 58 cats (16%) presented both MgE and abnormal CSF (five with only  
218 increased TNCC, two with both increased TNCC and total protein and two with  
219 albuminocytologic dissociation). MgE was detected in 17/58 cats (29%) with normal  
220 CSF. Abnormal CSF was seen in a total of 10/32 cats (31%) without MgE: 6 with only  
221 increased TNCC, three with both increased TNCC and total protein and one with  
222 albuminocytologic dissociation. No significant association ( $\chi^2$  test,  $P = 0.79$ ) was found  
223 between CSF results and MgE findings (Table I).

224 CSF abnormalities were more commonly detected in acute cases (16/37)  
225 compared to chronic cases (3/21), the difference being statistically significant (Fischer  
226 test,  $P = 0.04$ ), while MgE was similar in acute (15/37) and chronic (11/21) cases.  
227 Furthermore, none of the chronic cases presented an abnormal CSF without MgE (Table  
228 II). No association was found between the use of anti-inflammatory drugs before MRI  
229 and CSF analysis (Table III) or the presence or absence of a polyp (Table IV) and MgE  
230 or CSF abnormalities.

231

232 Bacterial culture was performed in 45/58 cases. Samples were collected from bulla  
233 osteotomy in 13 cases, myringotomy in 28 cases, and external ear canal in four cases.  
234 Negative bacterial culture was observed in 33/45 cases, of which thirteen (39%) received  
235 antimicrobial therapy before sampling. The percentage of negative bacterial culture was  
236 73% in both acute (21/30) and chronic (11/15) cases.

237 Twelve cases showed a positive bacterial culture: six for *Staphylococcus* species,  
238 three for *Pasteurella* species, one for *Streptococcus canis*, one for *Actinomyces pyogenes*  
239 and one with both *Streptococcus equi subspecies zooepidemicus* and *Staphylococcus felis*.  
240 Among these cases, seven out of twelve (58%) received antimicrobial therapy before  
241 sampling.

242 The results of the bacterial culture depending on presence/absence of MgE and  
243 normal/abnormal CSF findings are summarized in Table V. No statistical association was

244 found between a positive bacterial culture and MgE ( $\chi^2$ ,  $P = 0.82$ ) or CSF results ( $\chi^2$  test,  
245  $P = 0.15$ ). One observation was that if no MgE was seen on MRI images and no  
246 abnormality was detected on CSF analysis, the likelihood to get a **negative ear sample**  
247 bacterial culture from myringotomy or bulla osteotomy was only about 12% (2/17 cases).  
248 One cat had a positive culture from CSF (*Clostridium beijerinckii* and *Enterococcus*  
249 *faecalis*) despite no positive bacterial culture ear sample from myringotomy and  
250 clindamycin therapy for three days.

251

252         Twenty-five cats received corticosteroids and 13 received non-steroidal anti-  
253 inflammatory drugs after diagnosis. Choice of anti-inflammatory drugs according to  
254 MgE, CSF or bacteriology findings is summarized in Table VI. No significant difference  
255 ( $\chi^2$  test,  $P = 0.53$  for MgE; Fischer test,  $P = 0.08$  for CSF; Fischer test,  $P = 0.62$  for  
256 bacteriology) was identified although corticosteroids seem to be chosen more often in  
257 case of abnormal CSF (11 cases versus 1 case). A total of 54 cats (93%) received  
258 antimicrobial therapy after diagnosis. Twenty-six out of 54 (48%) had antimicrobial  
259 therapy started prior to referral while it was started by the referral centre after diagnosis  
260 in twenty-eight cases (52%). Thirty-nine received first line antimicrobials, seven received  
261 second line and eight received both. Duration of antimicrobial treatment depending on  
262 MgE, CSF or bacteriology findings are summarized in Table VII. Antimicrobial therapy  
263 duration tended to be longer in case of positive culture (5.58 vs 4.22 weeks) or when CSF



264 findings were abnormal (5.83 vs 4.76 weeks), although this difference was not  
265 statistically significant (Student's t-test).

266 Association between CSF results, MgE and outcome are described in table VIII.  
267 Good outcome is defined by an improvement of clinical signs (and euthanasia unrelated  
268 to the disease after several months). Poor outcome is defined by an absence of  
269 improvement or euthanasia. Although cases with MgE tend to have poorer outcome  
270 (5/18) than without MgE (3/26), the difference was not statistically relevant (Fischer test,  
271  $P = 0.24$ ). No association was found between outcome and CSF abnormalities (Fischer  
272 test,  $P = 1$ ). Bulla osteotomy was performed in 14 cases (8 polyps and 6 OMI), between  
273 one and 78 days after diagnosis (median = 5). Delayed bulla osteotomies were due to  
274 absence of improvement or relapse of clinical signs after initial medical management.  
275 Improvement of neurological signs after surgical management was seen in 11 cases (time  
276 of follow-up varies between one and 104 weeks including five cases with more than eight  
277 weeks of follow-up), relapse in one (three months after surgery) and three cases were lost  
278 to follow-up. None of the surgical cases were euthanized for reasons related to the OMI.  
279 Medical management resulted in improvement in 23/44 cases (time of follow-up varies  
280 between one and 78 months including 12 cases with more than eight weeks of follow-  
281 up). Two cases improved and were euthanized due to unrelated reasons four months and  
282 17 months after diagnosis of OMI respectively due to carcinoma and polyarthritis. Three  
283 cases did not improve after one month but owners declined surgery. Four cases were

284 euthanized following diagnosis or several weeks after with no improvement on medical  
285 therapy. One case showed intermittent vestibular signs and was euthanized 20 months  
286 after diagnosis due to seizure-like episodes. Eleven cases were lost to follow-up.

287 **Discussion**

288           The findings of this study show that in a cohort of 58 cats with PVS diagnosed  
289 with OMI, MgE is seen in approximately 50% of cases but only 27% of these cases had  
290 an abnormal CSF analysis. There was no association between MgE and abnormal CSF  
291 results. Chronic cases had significantly fewer abnormal CSF findings. When a treatment  
292 was given, its duration was similar regardless of a positive bacterial culture, abnormal  
293 CSF analysis or meningeal enhancement. Cases with MgE tend to have poorer outcome.

294

295           The use of MRI to investigate the cause of PVS is common practice in clinical  
296 neurology, particularly if concurrent meningitis is suspected. Administration of  
297 gadolinium-based contrast medium in MRI has been demonstrated as a more sensitive  
298 method to diagnose experimental bacterial meningitis in dogs compared to MRI without  
299 contrast or post-contrast CT (23). However, only severe meningeal inflammation at  
300 necropsy was correlated with MRI findings while mild inflammation was not detected in  
301 MRI leading to the conclusion that the absence of meningeal enhancement does not rule  
302 out bacterial meningitis (24). False positive results of diffuse MgE on MRI images  
303 without CSF abnormalities have also been previously described (25). MRI also allows a  
304 better evaluation of fluid containing inner ear structures than CT and otitis interna can be  
305 assessed on T2W images (16,26). More recently, d'Anjou *et al.* (2012) and Keenihan *et*  
306 *al.* (2013) compared MR sequences for detecting MgE in dogs. Post-contrast T1W and

307 T1W fat suppression were found to be the sequences of choice to detect meningeal  
308 inflammation (27,28). In previous studies in cats with OMI, only one case with focal  
309 meningitis resembled our population of cats with MgE, without intra-axial lesion or  
310 empyema. This case had an abnormal CSF analysis (1). In dogs, few studies describe  
311 naturally occurring OMI leading to meningeal enhancement in T1W images after  
312 gadolinium administration but no large study focusing on this aspect has been done;  
313 therefore its clinical consideration remains unknown (16,29,30).

314         Despite the first hypothesis that MgE is associated with abnormal CSF findings,  
315 a discrepancy between MgE and CSF results was found in 47% of cases. Indeed, MgE  
316 was seen more frequently in cases with unremarkable CSF analysis (65%) and abnormal  
317 CSF can be found without MgE in up to 31% of cases. To the best of the authors'  
318 knowledge, no previous report of cats with OMI evaluated the association between the  
319 MRI findings and CSF analysis. In previous reports of cats diagnosed with OMI and intra-  
320 axial lesions or empyema on MRI, an abnormal CSF was detected in 22/25 cases in which  
321 CSF analysis was performed (1,7,31). MgE was specifically described in one of these  
322 studies, where none of the five cats with chronic vestibular clinical signs had a MgE while  
323 all of the six acute or subacute cases showed MgE (7). The prevalence of MgE is in  
324 contrast with our results, in which 11 of 21 chronic cases and 15 of 35 acute cases showed  
325 MgE. When available, abnormal CSF analysis was detected in all acute or subacute cases  
326 (5/5) and only in 1 of 4 chronic cases (7). These results reflect our findings with

327 significantly fewer CSF abnormalities in chronic cases. Previous treatment with anti-  
328 inflammatory drugs or presence of a polyp did not affect presence/absence of MgE or  
329 CSF results.

330

331 The second hypothesis was that there would be an association between MgE  
332 and/or CSF abnormalities and a positive bacteriology culture. In case of absent MgE and  
333 normal CSF, the likelihood of a positive bacterial culture from myringotomy or bulla  
334 osteotomy is low (~ 12%). Even if this result is not statistically significant, it could make  
335 the clinician aware of the possible need to alter antimicrobial therapy after receiving  
336 culture results. Bacteriology results between acute or chronic cases did not differ  
337 significantly.

338 We finally hypothesized that positive bacterial culture would influence the choice  
339 and length of antimicrobial and/or anti-inflammatory therapy. Due to the retrospective  
340 and multicentre aspect of our study, medical management was very variable, making the  
341 investigation of the last hypothesis difficult. Generally, a long term (4 to 8 weeks), broad-  
342 spectrum antimicrobial therapy or, ideally, a therapy based on *in vitro* antimicrobial  
343 sensitivity profile, is recommended to treat OMI in cats and dogs (8). In the current study,  
344 duration of therapy was slightly longer in cases of positive bacterial culture than in cases  
345 of negative bacterial growth culture (respectively 5.58 weeks vs 4.22 weeks), although

346 the difference was not significant. The similar length of therapy may be due to initial  
347 prescription with no improvement of clinical signs or a lack of short-term follow-up by  
348 the referring centre. Also, if anti-inflammatory drugs were to be implemented after  
349 abnormal CSF results, clinicians tended to use corticosteroids more frequently. These  
350 results could reflect a clinicians' preference for corticosteroids in case of CNS  
351 inflammation. However, clinicians need to remember about the lack of CSF abnormalities  
352 in chronic cases, despite meningeal inflammation.

353 Culture results from samples taken from the external ear canal have to be  
354 interpreted with caution. Common microorganisms can be detected in the tympanic bulla  
355 of healthy cats in up to 25% of cases (32). Bacteria have been previously cultured from  
356 48% of healthy canine external ears (33). Moreover, up to 67% of myringotomies  
357 performed via video-otoscopy might be contaminated even if microorganisms were  
358 detected only in 15.4% of the samples (34). The presence or absence of bacteria on culture  
359 should not be considered as critical in formulating a treatment plan as the type of bacteria  
360 that are cultured (that is, whether they are likely of external ear canal origin and/or  
361 possible iatrogenic contaminants versus a likely cause of middle ear infection). The lack  
362 of a cultured infectious agent in our case series with the presumed presence of OMI raises  
363 the question of a purely inflammatory mechanism leading to otitis interna.

364 Age of cats, uni- or bilateral vestibular signs distribution, proportion of OMI with  
365 or without polyps and type of bacteria cultured were similar to previously published

366 literature (1,2,13,35). In this cohort cases with identified MgE tend to have poorer  
367 prognosis than those without MgE however this difference was not statistically  
368 significant, while an abnormal CSF result was not associated with any difference in  
369 outcome. This finding might help clinicians to anticipate and maybe adapt the therapy for  
370 such cases. Surgical treatment with bulla osteotomy was performed in fourteen cats  
371 (24%), including four in which it was performed several weeks after diagnosis. This was  
372 different to other published studies in which none of the cases with vestibular signs  
373 underwent surgery and up to 30% of otitis media cases without neurological signs  
374 received bulla osteotomy (1,4). In another study focusing on OMI in cats with intracranial  
375 complications, ventral bulla osteotomy was performed more often (12/18 cases, 67%)  
376 (31). Surgically managed cases lead to an improvement of neurological status for all cases  
377 (13/13) while medical management showed an improvement in 23/31 cases (74%). These  
378 results are slightly better than those for cats with OMI and intracranial complication (31).

379

380         There are several limitations to this study. First of all, the retrospective nature is  
381 associated with incomplete data and did not allow long term follow-up of cases. A  
382 multicentre study with different MRI machines, clinicians and protocols will result in  
383 differences in the evaluation of MR images and the clinical management of cases.  
384 Concerning case recruitment, cases without MRI and/or CSF analysis were excluded,  
385 which may have biased the study population towards potentially more severely affected

386 cases. Clinicians may decide against CSF analyses in those cases where clear MgE is seen  
387 on MRI, biasing the population towards a higher number of cases without MgE. A similar  
388 bias could also impact the choice of therapy in the cases of the present study. Moreover,  
389 the use of medication prior to presentation might have impacted the results of the current  
390 study. We decided to exclude all cases presented with intra-axial lesions or empyema on  
391 MRI images despite presumed peripheral vestibular lesion localisation as brainstem signs  
392 may not be clinically obvious in the neurological examination and might have impacted  
393 CSF results (36,37). In dogs, it has been shown that bacterial culture obtained via  
394 myringotomy can be contaminated by bacteria from the external ear canal and this could  
395 be one limitation for our bacteriology results (34). Time of acquisition in dogs and higher  
396 dose of gadolinium in humans may induce false negative meningeal enhancement  
397 (28,38). False positive MgE may also occur (25).

398



399 **Conclusion**

400 In this study, no association was found between MgE and CSF results. Nearly half  
401 of the cases (47%) showed a discrepancy between MRI and CSF findings. Additionally,  
402 the lack of MgE in MRI does not rule out the presence of a meningitis pathologically.  
403 Hence, CSF analysis may be useful to detect the presence of possible concurrent  
404 meningitis in cats with OMI. CSF findings and MgE results were not associated with  
405 likelihood of a positive or negative middle ear bacterial culture. Abnormal CSF results  
406 seemed to influence the clinicians' choice of anti-inflammatory drugs with a preference  
407 for glucocorticoids versus non-steroidal anti-inflammatory drugs. Abnormal CSF results  
408 were seen less frequently in chronic cases than acute cases in this study. Additionally, the  
409 identification of an abnormal CSF analysis did not seem to notably influence the length  
410 of antimicrobial therapy, which remains the mainstay for this presumed infectious  
411 disease. Outcome tend to be poorer when MgE is detected on MRI.

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523

524 **Figures**

525 Figure 1 - MRI (Philips Panorama HFO 1.0 T) transverse images in T1W sequence (a)  
526 and T1W post-contrast sequence (b) of a cat presented with an otitis media-interna  
527 without polyp (\*) and without meningeal enhancement.

528

529 Figure 2 - MRI (Philips Panorama HFO 1.0 T) transverse images in T1W sequence (a),  
530 T1W post-contrast sequence (b) and T2W sequence (c) of a cat presented with a bilateral  
531 otitis media-interna without polyp (\*), with meningeal and vestibulocochlear nerve  
532 enhancement (white arrowhead) and otitis interna (yellow arrowhead).

533

534 Figure 3 - MRI (Philips Panorama HFO 1.0 T) transverse images in T1W sequence (a)  
535 and T1W post-contrast sequence (b) of a cat presented with an otitis media-interna  
536 associated with a polyp (X) and without meningeal enhancement.

537

538 Figure 4 - MRI (Philips Panorama HFO 1.0 T) transverse images in T1W sequence (a)  
539 and T1W post-contrast sequence (b) of a cat presented with an otitis media-interna  
540 associated with a polyp (X) and with meningeal and vestibulocochlear nerve  
541 enhancement (white arrowhead).

542 **Supplementary material**

543 The following file is available online: table of cases

544

545

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556 **Ethical Approval**

557 The work described in this manuscript involved the use of non-experimental (owned or

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559 of veterinary clinical care for the individual patient were always followed and/or this work  
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563

564

565 **Informed consent**

566 Informed consent (verbal or written) was obtained from the owner or legal custodian of  
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