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Tuboimpedance – A New Test of Eustachian Tube Function

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2 **MANUSCRIPT**

3 **Tuboimpedance – A New Test of Eustachian Tube Function**

4

5 **Abstract**

6 Objective

7 Eustachian tube (ET) dysfunction is most frequently caused by a failure of the ET to
8 adequately open, however there is currently no reliable method of assessing this.
9 Tubomanometry has recently shown good inter-individual repeatability as a measure
10 of ET function, by measuring middle ear pressure after the application of regulated
11 nasopharyngeal pressures during swallowing. We present the first reports of a novel
12 test: middle ear impedance measurements during standard nasopharyngeal pressure
13 application (tuboimpedance). We assess repeatability in healthy ears, and any
14 advantages over tubomanometry.

15

16 Study Design

17 Exploratory cohort diagnosis study.

18

19 Setting

20 Tertiary referral center.

21

22 Subjects

23 20 screened, healthy ears (10 volunteers).

24

25 Methods

26 Tubomanometry and tuboimpedance tests were performed while individuals
27 swallowed with applied nasopharyngeal pressures of 20, 30, 40 and 50mbar.

28 Eustachian tube opening detection rate and test repeatability (measured by intraclass
29 correlation coefficient) for immediate and delayed repeats at each pressure were
30 compared.

31

32 Results

33 ET opening was detected more frequently using tuboimpedance, with a 100%
34 detection rate using a nasopharyngeal pressure of 30mbar or more, compared to 88-
35 96% with tubomanometry. Detection of ET opening at 20mbar was possible with
36 tuboimpedance. Repeatability of both tests was mostly strong (ICC >0.7) for both
37 immediate and delayed repeats. Repeatability for the tubomanometry R value was
38 only fair to moderate.

39

40 Conclusion

41 Tuboimpedance may provide a repeatable measure of ET opening that is easier to
42 perform, due to lower nasopharyngeal pressures required and fewer issues with poor
43 ear-probe sealing. Further assessment in patients with different forms of ET
44 dysfunction is required.

45

46 Word count 246

47

48 **Introduction**

49 Normal function of the Eustachian tube (ET) permits atmospheric equalisation of
50 middle ear pressure and mucociliary clearance of secretions, while protecting the
51 tympanic cavity from nasopharyngeal sounds and secretions. These processes are
52 facilitated through the intermittent and brief opening of the normally closed ET.
53 Despite a significant level of morbidity from Eustachian tube dysfunction (ETD)¹,
54 there has been no consensus on the optimal clinical test to detect ET opening, and
55 diagnosis is currently largely made on the basis of clinical history and examination ².

56

57 It is desirable to develop a simple and reliable test of ET function, in order to permit
58 objective diagnosis and quantification of ETD. In most cases ETD is obstructive in
59 nature, due to a reduced rate or absence of ET opening ¹. A large number of different
60 tests for ET opening have been described, each with its own strengths and weaknesses
61 ³. Many tests require a patient to generate a nasopharyngeal pressure, which is then
62 transmitted to the middle ear if the ET opens. Patients are taught to generate these
63 pressures by performing a manoeuvre such as a Valsalva (forcibly exhaling with the
64 nose and mouth occluded). However, the nasopharyngeal pressures generated by
65 individuals significantly vary, limiting the comparability of results between ears ⁴.

66

67 Tubomanometry has established itself as a test of ET function in recent years as it
68 enables the standardisation of nasopharyngeal pressures. The test is performed using
69 a device to automatically increase the nasopharyngeal pressure to typically 30, 40 or
70 50mbar via a sealed nosepiece. This pressure increase is timed to coincide with
71 patient swallowing. Tympanic pressure increases caused by ET opening are then
72 transmitted to the external auditory canal (EAC) via the mobile tympanic membrane,
73 and recorded with a sealed earpiece. By removing the need for the patient to perform

74 a Valsalva it not only ensures an adequate nasopharyngeal pressure is created, but it
75 allows the individual to swallow simultaneously. Tubomanometry therefore measures
76 both active ET opening (due to paratubal muscle activity with swallowing), and
77 passive forced opening due to the high positive pressure at the nasopharyngeal ostium
78 ⁵. This standardised, dual-assessment of both active and passive opening is not
79 possible if relying on patient manoeuvres alone, and tubomanometry has shown good
80 inter-individual repeatability ⁶.

81

82 Another method of detecting ET opening is with tympanic membrane impedance.
83 Our group has found that continuous impedance monitoring during various patient
84 manoeuvres provides a simple and repeatable method for detecting ET opening.
85 Unlike external auditory canal (EAC) manometry, the use of continuous impedance
86 assessment has been not been commonly used to assess ET opening ⁷.

87

88 We have investigated the feasibility of performing continuous impedance recording
89 using the tubomanometry method of inducing metered nasopharyngeal pressures in
90 time with swallowing, i.e. ‘tuboimpedance’. As a first investigation into this new
91 technique, the detected ET opening rate, and the repeatability of this method using
92 immediate and delayed repeats in healthy ears are compared to paired tubomanometry
93 results.

94

95 **Methods**

96 Independent ethical approval was obtained from the UK National Research Ethics
97 Service (South Cambridgeshire Committee).

98

99 *Candidates*

100 Volunteers without ear disease were recruited by advertisement and each volunteer
101 had both ears assessed independently. All volunteers underwent otoscopy, and those
102 with an abnormal tympanic membrane were excluded. In addition, volunteers were
103 required to score less than 14 on the 7-item Eustachian Tube Dysfunction
104 Questionnaire (ETDQ-7) ⁸, and have bilateral Jerger Type A tympanograms ⁹ (226Hz
105 tone at 85dB SPL, Titan IMP440, Interacoustics, Assens, Denmark).

106

107 *Equipment*

108 A tubomanometry device (Tubomanometer, Spiggle & Theis, Overath, Germany) was
109 used to generate nasopharyngeal pressures via a two-pronged nosepiece fitted to both
110 nostrils. To start the testing the tubomanometer EAC pressure sensor was sealed into
111 the right ear canal. A probe producing a 226Hz 85dB SPL tone was sealed in the left
112 ear canal for continuous impedance recording (JK-05AD, Rion Co., Tokyo, Japan).
113 The JK-05AD was also connected to the nasal circuit to allow the nasopharyngeal
114 pressure to be displayed alongside the impedance trace. The apparatus is illustrated
115 schematically in Figure 1.

116

117 *Data collection and interpretation*

118 Volunteers were requested to swallow a small water bolus, automatically triggering
119 the nasopharyngeal pressure increase. Measurements from both ears were recorded
120 simultaneously. Nasopharyngeal pressures selected for investigation were 30, 40 and
121 50mbar, according to current standard practice ⁶. In addition 20mbar was trialled to
122 assess if a lower pressure might be adequate for testing, while being better tolerated
123 by the volunteers. Nasopharyngeal pressures were applied in order of increasing
124 magnitude, and testing at each pressure was repeated twice (immediate repeat data).
125 After a set of results was collected, the tubomanometry and impedance earpieces were

126 swapped, and a further data-set of two repeats at each pressure was recorded from the
127 opposite ears. The complete process was then performed again after an interval of
128 around 15 minutes, to record delayed values for each of the tests in both ears (delayed
129 repeat data).

130

131 Impedance was measured in units of equivalent volume of air in ml ¹⁰, and a positive
132 deflection from baseline ≥ 0.05 ml was considered positive for ET opening ⁷. For
133 tubomanometry, a positive EAC pressure increase greater than 0.1mbar was recorded
134 as an opening. Based on the shape of the pressure or impedance traces, recordings
135 were assigned to one of two groups: persistent or non-persistent middle ear pressure.
136 Volunteers were requested not to swallow immediately following the test, and if the
137 impedance or EAC pressure trace was maintained at greater than 50% of the peak
138 value at one second after nasopharyngeal pressure returned to normal, the middle ear
139 pressure was recorded as persistent (Figure 2). As has become standard practice, the
140 R-value, a measure of the latency of the middle ear pressure change (opening) with
141 respect to the nasopharyngeal pressure was also calculated ⁵. Early opening of the ET
142 ($R \leq 1$) is thought to indicate normal ET function, and late opening ($R > 1$) to suggest
143 impaired ET opening ⁶.

144

145 Data were analysed at the single-ear level using Microsoft Excel and IBM SPSS. Test
146 results were assessed for the repeatability of both immediate and delayed repeats with
147 the intraclass correlation coefficient (ICC), calculated with a mixed effects model
148 assessing absolute agreement. The ICC is measured on a scale of 0 to 1, where 1
149 represents perfect reliability with no measurement error, and 0 indicates no reliability.

150

151 **Results**

152 Twenty healthy ears from ten volunteers (five male, mean age 22) were recruited.
153 The mean ETDQ-7 score was nine (range 7-12). All volunteers were able to complete
154 the assessment in full.

155

156 Results from tuboimpedance and tubomanometry testing are presented in Table 1. ET
157 opening was detected more frequently with the tuboimpedance method, with a 100%
158 detection rate using a nasopharyngeal pressure of 30mbar or more. Between 57% and
159 88% of middle ear pressure changes were classed as persistent.

160

161 Repeatability of the tests for equivalent volume values (tuboimpedance) and EAC
162 pressure values (tubomanometry) was very good for both immediate and delayed
163 repeats, as measured by ICC. Where the mean of two immediate repeat values was
164 used, the ICC values obtained when comparing the initial and delayed measurements
165 was further improved. Immediate and delayed repeatability for the tubomanometry
166 R-value was variable between 0.2 (poor) and 0.83 (almost perfect), and overall R-
167 value repeatability was poorer than that of EAC pressure.

168

169 **Discussion**

170 This study demonstrates the feasibility of performing tuboimpedance, a hybrid of the
171 increasingly popular tubomanometry test and the continuous impedance test.

172

173 At all pressures tested, tuboimpedance detected more openings than tubomanometry,
174 with a maximum difference of 12% at 30mbar. As our results for the two tests came
175 from a single cohort of ears, this difference suggests that tubomanometry fails to
176 detect some openings that can be detected by tuboimpedance. The reason for this
177 finding is not clear, but it may be that tuboimpedance can detect subtle changes in TM

178 stiffness that do not translate to a TM movement, and therefore EAC pressure does
179 not change. Despite the small numbers, our tubomanometry data are comparable to
180 published data, reinforcing this finding: In our cohort of healthy ears we were unable
181 to measure ET opening in a mean of 8% of test repeats across the 30-50mbar
182 pressures, while Esteve *et al.* and Schroder *et al.* reported rates of 7% ⁵ and 4% ⁶
183 respectively.

184

185 We also found that the tuboimpedance test is better at detecting ET opening than
186 standard continuous impedance testing that relies on patient manoeuvres to generate
187 pressures. In initial work from our group using a similar healthy cohort, impedance
188 testing detected passive ET opening in 88% of Valsalva manoeuvres. Only one other
189 published account of the continuous impedance technique for ET testing could be
190 found, with a Valsalva-associated ET opening rate of 93% in healthy ears ⁷. In
191 comparison, in our tuboimpedance data, using the standard tubomanometry pressures
192 of 30, 40 and 50mbar, 100% of tests resulted in detection of ET opening. This
193 increase in opening detection over Valsalva testing is likely due to the combined
194 active and passive ET opening occurring with both paratubal muscle contractions, and
195 a large positive pressure differential across the ET. It is desirable to test active, as
196 well as passive opening when assessing ET function, as failures of either action alone
197 may lead to ETD ².

198

199 Most middle ear pressure changes appeared to persist beyond the applied
200 nasopharyngeal pressure, with similar findings in both tests. The persistence is due to
201 trapping of pressure within the middle ear, in the absence of swallowing to actively
202 open the ET. Typically, even in traces classed as persistent, there was some loss of
203 middle ear pressure from escape along the ET. It is hypothesised that persistence was

204 less frequent at higher nasopharyngeal pressures as the larger induced middle ear
205 pressures were more likely to passively force the ET open and leave a residual
206 pressure <50% of the peak pressure. The magnitude of residual middle ear pressure
207 both before after further swallows may provide additional diagnostic information in
208 ETD cases.

209

210 The rationale behind using three different pressures in tubomanometry is that it allows
211 quantification of ETD, with individuals experiencing less severe ETD demonstrating
212 opening and normalisation of R-values at higher applied pressures⁵. Through clinical
213 use, we have found that some individuals struggle to perform tubomanometry using
214 the highest (50mbar) pressure, as it can be difficult to obtain a seal for the nosepiece,
215 and some do not tolerate the sudden large pressure increase. While the range of
216 nasopharyngeal test pressures is a useful feature of the test, it is desirable for these to
217 be as low as possible, to ease test performance and reduce patient discomfort.

218

219 At the non-standard, lower pressure of 20mbar, tubomanometry detected opening with
220 86% of swallows. However, the opening detection rate was better maintained with
221 tuboimpedance, which at 91% is more similar to rates seen at 30mbar in
222 tubomanometry^{5,6}. It may be that tuboimpedance could routinely be used with 20, 30
223 and 40mbar pressures, maintaining the ability to quantify ETD severity, but making it
224 easier to perform and more feasible to use routinely in a clinical setting. This would
225 be of particular use for those who might struggle with 50mbar pressures, such as
226 children and the elderly.

227

228 Tuboimpedance has been shown to have good repeatability. With both immediate
229 and delayed repeat testing, the consistency of equivalent volume and EAC pressure

230 values for tuboimpedance and tubomanometry respectively were very good, as
231 measured by the ICC. For both tests, by taking the mean of two immediate test
232 repetitions, the repeatability of results over a 15 minute delay can be improved.
233 Performing each test more than once and using the mean value may therefore have a
234 role in clinical use, particularly given the inherent variation found with each swallow.

235

236 The R-value is a derived value used in tubomanometry to reduce inter-subject
237 variability, and classify ears with ET opening as either $R \leq 1$ (normal), or $R > 1$ (mild
238 ETD). A similar value could be derived from tuboimpedance traces if desired,
239 however we found the repeatability for the tubomanometry R-value to be worse than
240 that of the pressure values. When used simply to classify ears as $R \leq 1$ or $R > 1$
241 Schroder *et al.* reported ICC values of 0.83-0.90⁶, but our comparable ICC values
242 were 0.48-0.52 for immediate repeats and 0.62-0.69 for delayed repeats. The reason
243 for this difference is not clear.

244

245 An advantage of tuboimpedance over tubomanometry was that fitting the impedance
246 earpiece was straightforward and reliable, as although the earpiece requires a secure
247 fit, it does not need to be hermetically sealed. In contrast, maintaining an air-tight
248 seal in the external auditory canal for pressure tubomanometry measurements proved
249 challenging.

250

251 The main drawback of tuboimpedance is that it cannot be performed if the tympanic
252 membrane is perforated or has a grommet in situ, whereas tubomanometry can still be
253 performed by adjusting the scale on the EAC pressure display. Both techniques will
254 fail to reliably record openings in the presence of a middle ear effusion.

255

256 Swallowing causes reflex contraction of both the paratubal muscles and the tensor
257 tympani muscle ¹¹. Tensor tympani contraction has been shown under normal
258 auditory canal pressures to increase tympanic impedance during swallowing ¹², and
259 there is therefore the potential for false positive impedance findings during swallows.
260 However, tensor tympani contracts for approximately 300ms during swallowing ¹³,
261 whereas the majority of traces at each pressure demonstrated persistence of the
262 impedance change after the nasopharyngeal pressure had returned to normal, and long
263 after the initiating swallow. False positive findings due to tensor tympani contraction
264 are therefore not thought to be significant.

265

266 The thresholds used to assign recordings as openings were set based on published
267 literature and past experience with the devices. The lack of a reference standard for
268 detecting Eustachian tube opening prevents confirmation that these thresholds are
269 optimised to reduce false negatives or positives, and further experience with the
270 techniques in ETD cases is required to refine this.

271

272 **Conclusion**

273 ETD is a common condition where a lack of diagnostic tools and outcome measures
274 has hindered clinical practice and research into new treatments. Tuboimpedance is a
275 novel hybrid test, with our pilot testing indicating that it may be superior to
276 tubomanometry in its ability to detect ET opening in healthy ears, as well as being
277 easier to use. Tuboimpedance is at an early stage of development, but should now be
278 trialled in adult and paediatric patients with ETD to assess whether diagnostic
279 accuracy can match or improve upon that measured with tubomanometry.

280

281 **Acknowledgments**

282 None

283

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320 man during swallowing. *Acta oto-laryngologica*. May-Jun 1978;85(5-
321 6):453-455.
- 322
- 323
- 324

325 **Table 1.** Detected Eustachian tube opening rate and repeatability with tuboimpedance
 326 and tubomanometry

| Test | Naso-pharyngeal pressure | % opening detected | % ME pressure persistent | Immediate repeat ICC | Delayed repeat ICC |
|---------------------------------|---------------------------------|---------------------------|---------------------------------|-----------------------------|---------------------------|
| Tuboimpedance Equivalent Volume | 20mbar | 91 | 88 | 0.75 | 0.70 |
| | 30mbar | 100 | 78 | 0.71 | 0.61 |
| | 40mbar | 100 | 68 | 0.85 | 0.80 |
| | 50mbar | 100 | 60 | 0.82 | 0.72 |
| Tubomanometry EAC Pressure | 20mbar | 86 | 78 | 0.77 | 0.72 |
| | 30mbar | 88 | 75 | 0.77 | 0.79 |
| | 40mbar | 93 | 63 | 0.72 | 0.71 |
| | 50mbar | 96 | 57 | 0.73 | 0.46 |
| Tubomanometry R-Value | 20mbar | NA | NA | 0.32 | 0.66 |
| | 30mbar | | | 0.65 | 0.61 |
| | 40mbar | | | 0.46 | 0.39 |
| | 50mbar | | | 0.20 | 0.28 |

327

328

329 **Figure / Table Legends**

330

331 Figure 1.

332 The Tubomanometer was connected in the usual way, with probes sealed in the EAC
333 and nose. The impedance probe was sealed in the contralateral ear, and a feed taken
334 from the nosepiece so that nasopharyngeal pressure could be displayed on both
335 machines.

336

337 Figure 2.

338 Diagram representation of persistent and non-persistent impedance or EAC pressure
339 traces. If the trace height was >50% of the peak height 1 second after the
340 nasopharyngeal pressure returned to normal the middle ear pressure was recorded as
341 persistent.

342

343 Table 1.

344 Percentage of detected openings with swallows and intraclass correlation coefficient
345 (ICC) for tuboimpedance and tubomanometry repeats. Immediate repeats were
346 performed consecutively and delayed repeats after approximately 15 minutes. The
347 mean values of the immediate repeats were also compared. Intraclass Correlation
348 Coefficient (ICC) can be interpreted as follows: 0-0.2 poor agreement; 0.3-0.4 fair
349 agreement; 0.5-0.6 moderate agreement; 0.7-0.8 strong agreement; and >0.8 almost
350 perfect agreement. EAC – External auditory canal.