

1 **3D imaging based web application for tracheal tube depth in preterm**
2 **neonates.**

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4 Raksa Tupprasoot MBBS¹, Dean Langan PhD², J Ciaran Hutchinson MBBS^{1,2}, Hannah
5 Barrett MBBS^{1,2}, Mike Sury MBBS^{1,2}, Owen Arthurs PhD FRCR^{1,2*}

6
7 **Affiliations:**

8 ¹ Great Ormond Street Hospital for Children NHS Foundation Trust, London, UK

9 ² UCL Great Ormond Street Institute of Child Health, University College London, London,
10 UK

11
12 **Address correspondence to:**

13 Dr Owen J Arthurs, Consultant Radiologist,

14 Great Ormond Street Hospital for Children NHS Foundation Trust, London, WC1N 3JH, UK

15 E mail: owen.arthurs@gosh.nhs.uk, Tel/Fax +44(0)20 7405 9200

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17 **Running header:** Tracheal tube depth application

18 **Key words**

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31

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33 The authors have no conflicts of interest relevant to this article to disclose.

34

35 What's known on this subject

36 Correct depth insertion of a tracheal tube (TT) is challenging in preterm infants. Currently
37 there is no reliable single-predictor model for neonates applicable to the whole range of size
38 or age.

39

40 **What this study Adds**

41 We used 3D fetal images to measure mid-tracheal length, to help predict ideal tracheal tube
42 insertion depth in preterm infants. Our best model is available as an easy to use internet
43 application, using 4 clinical variables.

44

45 **Contributors statement**

46

47 **Raksa Tupprasoot:** Dr Tupprasoot helped co-design the study, performed the literature
48 search, carried out the data analysis, and drafted the intial manuscript.

49 **Dean Langan:** Dr Langan performed the statistical analysis for the study, and critically
50 reviewed the manuscript.

51 **J Ciaran Hutchinson:** Dr Hutchinson carried out the data analysis and critically reviewed
52 the manuscript

53 **Hannah Barrett:** Dr Barrett carried out the data analysis and critically reviewed the
54 manuscript.

55 **Mike Sury:** Dr Sury co-designed the study, performed the literature search, and critically
56 reviewed the manuscript.

57 **Owen Arthurs:** Dr Arthurs co-designed the study, supervised the data collection, and
58 critically reviewed the manuscript.

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60 All authors have participated sufficiently in this submission and take public responsibility for
61 its content. All authors have approved the final version as submitted, and agree to be
62 accountable for all aspects of the work.

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65 Abbreviations:

66	PM MRI	post mortem magnetic resonance imaging
67	TT	Tracheal tube
68	TD	internal tracheal diameter
69	GA	gestational age
70	FL	foot length
71	CRL	crown-rump length
72	BW	body weight
73	3D	Three dimensional
74	Mid-TL	mid-tracheal length, defined as the distance between the lips and the
75		mid tracheal point
76	uAL	upper airway length, defined as the distance from the lips to the glottis.
77	total	total airway length, defined as the distance from the lips to the carina
78	TL	tracheal length = totAL-uAL

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82

83 **Abstract**

84 **Background:** Positioning a tracheal tube (TT) to the correct depth in pre-term infants is
85 challenging. Currently there is no reliable single-predictor model for neonates applicable to
86 the whole range of size or age.

87 **Objective:** In this study, we used post mortem magnetic resonance images from preterm
88 infants to measure tracheal dimensions and to develop a clinical guide for TT positioning.

89 **Methods:** We measured tracheal length and diameter in a cohort of normal neonates and
90 fetuses who underwent post mortem MRI (cause of death unexplained). The distance
91 between the lips and the mid tracheal point (mid-tracheal length = mid-TL) and tracheal
92 diameter (TD) was obtained. We produced univariate prediction models of mid-TL and TD,
93 using gestational age (GA), foot length (FL), crown-rump length (CRL) and body weight
94 (BW) as potential predictors, as well as multiple prediction models for mid-TL.

95 **Results:** Tracheal measurements were performed in 117 cases, with mean GA 28.8 w (range
96 14 to 42 w). The best linear relationship was between mid-TL and FL (mid-TL = FL * 0.914
97 + 1.859; $R^2=0.94$) but was improved by multivariate regression models. We developed a
98 prediction tool using only gestation and body weight ($R^2=0.92$) which is now available as a
99 web-based application via the internet.

100 **Conclusion:** Post mortem imaging data provides estimates of TT insertion depth. Our
101 prediction tool based on age and body weight can be used at the bedside and is ready to be
102 tested in clinical practice.

103

104 **Introduction**

105 Correct depth insertion of a tracheal tube (TT) is essential to avoid misplacement into the
106 bronchus or the pharynx, and this becomes more challenging as infant size decreases. Ideally,
107 the TT tip should be placed at the mid-point between the larynx and the carina, and although
108 its position can be checked by chest X-rays [1], repositioning is frequently necessary [2].

109

110 Methods used to investigate the correct or ideal TT depth have involved either imaging with
111 conventional chest radiographs or post mortem (PM) autopsy [3 – 6] and several formulae or
112 rules have been published to help accurately predict safe insertion depth. Studies have shown
113 that airway length and tube insertion depth have linear relationships with body weight [7],
114 gestation [8], foot length [9], and body size such as crown-rump [10] or crown-heel lengths
115 [4, 10]. The European Resuscitation Council has recommended that the TT depth estimation
116 should be based on gestation [11] although in practice, the difference between body weight
117 and gestation may not be appreciable [12]. However, most of the published studies have
118 involved too few very (<32-28 w) or extremely preterm (<28w) infants and relationships
119 change or become non-linear when infants less than 1 kg are included [8, 13, 14]. Currently
120 there is no reliable single-predictor model for neonates applicable to the whole range of size
121 or age.

122

123 Modern three dimensional (3D) cross-sectional imaging can be used to measure airway and
124 tracheal dimensions and should be more accurate than simple 2D chest radiography. 3D
125 imaging of airway structures is only rarely indicated in live preterm infants, but recently PM
126 magnetic resonance imaging (PMMRI) is being used routinely to investigate the cause of death

127 [15]. Our institutional autopsy imaging database provides 3D data on the airway dimensions
128 in a wide range of fetuses and neonates and could prove useful to develop a mathematical
129 model for the bedside. Furthermore our database includes fetuses younger than 22w gestation
130 who although being too preterm to survive, may be of future interest.

131

132 The aim of this study was to use 3D detailed anatomy derived from fetal PMMRI to measure
133 airway parameters, and to develop a bedside mathematical tool to predict optimal TT
134 insertion depth.

135

136 **Methods**

137 **Recruitment and criteria**

138 We evaluated PMMRI of all fetuses (miscarriages and stillbirths) aged less than 44 weeks
139 gestation referred to our institution from February 2012 to September 2015. Ethical approval
140 was obtained for analysis of PMMRI and written informed consent was obtained from
141 parents. Bodies were stored in a mortuary at 4°C until PMMRI. Cases were excluded if the
142 airway was abnormal on either PMMRI or subsequent autopsy. or where image quality was
143 inadequate to permit measurements. Demographic data acquired from the clinical notes
144 included gestational age (GA; weeks), body weight (BW; kg), foot length (FL; cm), and
145 crown-rump length (CRL; cm).

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149 **Magnetic Resonance Imaging**

150 Imaging was performed on a 1.5 T scanner (Avanto, Siemens Medical Solutions, Erlangen,
151 Germany) with a conventional phased array head coil. Conventional 3D T₁-weighted and T₂-
152 weighted sequences were examined by a pediatric radiologist for clinical purposes [16]. T₂-
153 weighted isotropic sequences of the head and chest were used to create 3D multi-planar
154 (sagittal, coronal and axial) datasets.

155

156 **Tracheal measurements**

157 Reformatted images (Figure 1), using a Centricity Web DX Viewer (Centricity WebPACS
158 system, 2006; GE Healthcare, Chalfont St Giles, UK) were used to measure and calculate the
159 following:

- 160 1. upper airway length (uAL) = distance from lips to glottis. The position of the glottis
161 was defined that part of the airway at the level of C5/C6 intervertebral disc space
162 because this has a close relationship with the cricoid cartilage;
- 163 2. total airway length (totAL) = distance from the lips to carina;
- 164 3. tracheal length (TL) = totAL-uAL;
- 165 4. the mid-tracheal length (mid-TL) = the distance between the lips and the mid-tracheal
166 point, and calculated as $uAL + \frac{1}{2}TL$; this is equivalent to a tracheal tube depth
- 167 5. internal luminal tracheal diameter (TD) measured at the mid-tracheal point.

168 All measurements were made to the nearest mm by a single observer (RS). Twenty datasets
169 were selected at random and measurements were repeated by a second observer, (OJA), to
170 assess inter-observer variability.

171

172 **Statistical analysis**

173 Univariate linear regression models were fitted for both outcome variables (mid-TL and TD)
174 using 4 predictors (GA, BW, FL and CRL). Two multivariate regression models were fitted
175 for mid-TL using (1) the two most readily available predictors (GA and BW and (2) all four
176 predictors. These prediction models were developed into a web application to for clinical
177 practice. For each regression model, subjects were identified in whom the model would have
178 predicted a mid-TL that would have resulted in a TT inserted either too short or too long (i.e.
179 the TT tip would be above the glottis or below the carina). Bland-Altman limits of agreement
180 were calculated to describe inter-observer variability of mid-TL and, using a regression
181 approach [17], to account for a relationship between variability and the mid-TL itself. All
182 analyses were carried out in R (version 3.3.0).

183

184

185 **Results**

186 Tracheal measurements were performed in 117 fetuses (mean GA 28.8 w, range 14 to 42 w;
187 17 fetuses were below 22 w, Table 1). The smallest infant weighed only 50g, and had a CRL
188 of 10cm. Mid-TL ranged between 2.8 and 10.8cm (Table 1).

189

190 All predictor variables had a strong linear relationship with mid-TL. FL had the highest
191 adjusted R^2 of 0.94 (Table 2) and produced the fewest predictions of tracheal tube tip
192 positioning below the carina 3 (2.6%) or above the glottis 2 (1.7%; Table 2 & Figure 2). BW
193 had the lowest adjusted R^2 of 0.86 but our results suggested that this may be because of a
194 non-linear relationship, particularly at low birth weights (log transformation R^2 0.91; Figure
195 2). The multivariate regression model using all four predictors had only a marginally better fit
196 than the multivariate model with only GA and BW (adjusted R^2 0.94 and 0.92 respectively;
197 Table 3).

198

199 Formulae for these models were made accessible through a web-based application
200 (<https://chpredict.shinyapps.io/shinyapp/> ; Figure 3).

201

202 TD was only measurable in 58 (50%) of fetuses. Univariate prediction models for TD all had
203 adjusted $R^2 = 0.51$ to 0.53 and multivariate regression modelling was not undertaken.

204 Variability (agreement between observers) of mid-TL increased as mid-TL increased: 95%
205 limits of agreement were ± 0.25 cm and ± 0.75 cm for mid-TL 4cm and 10cm respectively.

206

207 Discussion

208 We used fetal PMMRI 3D images to measure mid-tracheal length, in order to produce a
209 mathematical model to help predict the ideal tracheal tube insertion depth in preterm infants.
210 Our best model to predict mid-TL uses 4 clinical variables, but a model using only GA and
211 BW was almost as good. Tracheal diameter was not easily or accurately measured due to
212 small size.

213

214 Other investigators have used PM fetuses and neonates to measure ideal TT insertion depth.
215 Embleton and colleagues (2001) dissected 39 specimens ranging from 24 to 43 weeks post-
216 menstrual age and showed that FL was a much better predictor of TT depth ($R^2 = 0.79$)
217 compared to BW ($R^2 = 0.67$) and age ($R^2 = 0.58$) [9]. Neonatal body dimensions however,
218 such as foot length and crown rump length, are neither routinely measured at birth nor readily
219 achievable in an emergency intubation setting. A prediction model combining body
220 dimensions with BW and GA may be more slightly more accurate but is less practical in a
221 clinical situation than a model using GA and BW alone.

222

223 Previous studies in live infants have developed formulae based on age and weight. The 7-8-9
224 rule used BW to estimate TT insertion depth defined as the distance from the lips to the level
225 of the first or second thoracic vertebra on a chest radiograph [7]. The derived formula was
226 $\text{length} = 1.17 \times \text{BW} + 5.58$, which approximates to 6 + each kg body weight: this produces a
227 TT depth of 7 cm for 1 kg, 8 cm for 2 kg and 9 cm for 3 kg infants. The data in this study
228 from infants <1kg however were sparse, and Peterson and colleagues reported that the
229 formula gave TT depths that were too long in preterm infants <750g [13]. An internet tool

230 (currently available at <http://www.nicutools.org/>) uses the formula TT depth (cm) = 1.1 x BW
231 + 6.1, but only for infants >1 kg: for smaller infants the TT depth is 5.5 cm if <500g, 6 cm if
232 550 to 700g and 6.5 cm if 700 to 999 g. Kempley and colleagues reported that TT depth was
233 not linearly related to BW and that estimates based on GA reduced the need for TT
234 repositioning [8]. We found also that GA was not linearly related to mid-TL especially in our
235 smallest fetuses.

236

237 Nevertheless, a clinical study randomising neonates to receive a TT depth based on either GA
238 or BW suggested that there was no appreciable difference [12] and that neither predictor was
239 reliable at achieving satisfactory positioning: BW (the 7-8-9 rule) was successful in only 25
240 of 49 (51%) infants and GA was successful in 16 of 41 (39%) [18].

241

242 In light of these findings, our data and model may help to better predict the TT depth. Firstly,
243 our data is based on 3D anatomy of tracheal and airway measurements from MR imaging,
244 rather than two dimensional radiographic imaging using vertebral body heights as reference
245 levels for the trachea. Secondly, we provide new high quality data in the <22 week group
246 which increases the confidence in the mathematical model to predict mid-TL for potentially
247 viable infants of 23 to 25 week GA. Thirdly, our data supports the clinical findings of others
248 that any single predictor of TT depth is not as reliable as a combination of predictors.
249 Fourthly, by incorporating all our data, we have made available a web-based application,
250 which may be useful at the bedside. Whether using the data in this study improves ETT
251 placement accuracy remains to be determined in the appropriate clinical setting.

252

253 The main limitation of our study is that we did not measure the effect of the position of the
254 head and neck. Neck extension is known to lengthen the trachea [19 - 21] and imaging in a
255 defined neutral position would provide the most reliable predictions. There are physiological
256 changes which occur after death which may mean that our measurements will be different to
257 those in live infants. The trachea may be shorter at PM because the diaphragm applies less
258 traction [22] and therefore our formula may under-estimate mid-TL and TT insertion depth
259 for live infants. Collapse of the upper airway in a dead infant may account for a small degree
260 of measurement error and was most evident when we attempted to measure TD. Nevertheless
261 inter-observer variation was small and our measurements were repeatable. Our TT insertion
262 depths were also made to the nearest mm but clinicians may not be able to achieve accuracy
263 of insertion depth more than to the nearest 0.5cm; we recommend rounding up or down
264 appropriately. We look forward to [testing our formula in clinical practice](#) and potentially
265 improving it with additional PM imaging and clinical data.

266

267 **Conclusion**

268 PM imaging data provides reproducible anatomical measures of tracheal length in order to
269 predict ideal tracheal tube insertion depth. We have provided an easy to use internet
270 application which may be used at the bedside to improve TT tube placement. [This tool](#)
271 [remains to be validated in clinical practice.](#)

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336 **Table legends**

337

338 **Table 1.**

339 Summary of demographic details.

340 $\text{midTL} = \text{uAL} + \frac{1}{2} (\text{totAL} - \text{uAL})$

341

342 **Table 2.**

343 Univariate linear models of mid-TL

344

345 **Table 3.**

346 Multivariate linear models of mid-TL

347

348

349

350 **Figure legends**

351

352 **Figure 1: Airway measurements**

353 Example of multi-planar reconstruction (MPR) of PMMR sequence and tracheal
354 measurements – from mouth to carina (left and centre, top & bottom row), mouth to epiglottis
355 (right, top and bottom row)

356

357 **Figure 2. Relationship between mid-TL and GA**

358 Scatter plots of mid-TL against the four predictor variables; GA (top-left), FL (top-right),
359 CRL (bottom-left) and PMW (bottom-right). Regression lines (from table 2) are plotted in
360 red. Vertical lines represent absolute tracheal length (TL) in each case, and those in red
361 represent where predicted mid-TL falls outside this range.

362

363 **Figure 3. Screenshot of web-based application**

364 Formulae for both multiple prediction models of airway to mid tracheal length are currently
365 accessible through a web-based application situated at
366 <https://chpredict.shinyapps.io/shinyapp/>