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Title:

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Date:

2013-06-01

Citation:

Schmölzer, G. M., O'Reilly, M., Davis, P. G., Cheung, P. Y. & Roehr, C. C. (2013). Confirmation of correct tracheal tube placement in newborn infants. *Resuscitation*, 84 (6), pp.731-737. <https://doi.org/10.1016/j.resuscitation.2012.11.028>.

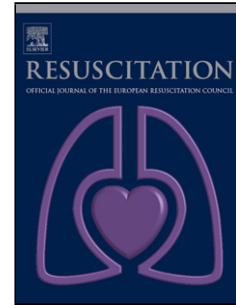
Persistent Link:

<https://hdl.handle.net/11343/43988>

Accepted Manuscript

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PII: S0300-9572(12)00930-6
DOI: doi:10.1016/j.resuscitation.2012.11.028
Reference: RESUS 5433

To appear in: *Resuscitation*

Received date: 21-7-2012
Revised date: 26-11-2012
Accepted date: 26-11-2012

Please cite this article as: Schmölzer GM, O'Reilly M, Davis PG, Cheung P-Y, Roehr CC, Confirmation of Correct Tracheal Tube Placement in Newborn Infants, *Resuscitation* (2010), doi:10.1016/j.resuscitation.2012.11.028

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1 **Confirmation of Correct Tracheal Tube Placement in Newborn Infants**

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24

25 **Abstract:** 139 words

26 **Word count:** 3821 words

27 **Appendix:** 1

28 **Figures:** 1

29 **Tables:** 2

30

31 **Short title:** Challenges of Endotracheal intubation in Newborns

32

33 **Conflict of interest:** None

34

35 **No reprints requested**

36

37 **Keywords:** INFANTS, NEWBORN, ENDOTRACHEAL INTUBATION, DELIVERY
38 ROOM, NEONATAL RESUSCITATION, NEONATAL INTENSIVE CARE UNIT

39

40 **Authors' affiliations**

41 GMS is supported by a Banting Postdoctoral Fellowship, Canadian Institute of
42 Health Research and an Alberta Innovate – Health Solution Clinical Fellowship. PGD
43 is supported by an Australian National Health and Medical Research Council
44 Practitioner and Principal Research Fellowship, respectively. PGD holds an
45 Australian National Health and Medical Research Council Program Grant No.
46 384100. PYC is an investigator of the Canadian Institutes of Health Research and
47 the Alberta Heritage Foundation for Medical Research. CCR is the recipient of a
48 European Respiratory Society Fellowship (LTRF fellowship n°15-2011).

49

50 **Author's contribution:**

51 Georg M. Schmölzer: Dr. Schmölzer conceptualized and designed the study, carried
52 out the initial literature search and analysis of available literature, drafted the initial
53 manuscript, and approved the final manuscript as submitted.

54

55 Megan O'Reilly: Dr. O'Reilly carried out the initial literature search and analysis of
56 available literature, reviewed and revised the manuscript, and approved the final
57 manuscript as submitted.

58

59 Peter G. Davis: Dr. Davis performed a literature search and analysis of available
60 literature, reviewed and revised the manuscript, and approved the final manuscript as
61 submitted.

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63 Po-Yin Cheung: Dr. Cheung performed a literature search and analysis of available
64 literature, reviewed and revised the manuscript, and approved the final manuscript as
65 submitted.

66

67 Charles Christoph Roehr: Dr. Roehr conceptualized and designed the study,
68 performed a literature search and analysis of available literature, reviewed and
69 revised the manuscript, and approved the final manuscript as submitted.

70

71 **Acknowledgement:**

72 We acknowledge the support of the European Respiratory Society, Fellowship LTRF
73 *fellowship n°15-2011* for Charles C Roehr.

74

75 **Abbreviations:**

76 NICU - Neonatal intensive care unit

77 DR - Delivery room

78 CO₂ - Carbon dioxide

79

80

Accepted Manuscript

80 **Abstract**

81 Tracheal intubation remains a common procedure during neonatal intensive care.
82 Rapid confirmation of correct tube placement is important because tube malposition
83 is associated with serious adverse outcomes. The current gold standard test to
84 confirm tube position is a chest radiograph, however this is often delayed until after
85 ventilation has commenced. Hence, point of care methods to confirm correct tube
86 placement have been developed. The aim of this article is to review the available
87 literature on tube placement in newborn infants. We reviewed books, resuscitation
88 manuals and articles from 1830 to the present with the search terms “Infant,
89 Newborn”, “Endotracheal intubation”, “Resuscitation”, “Clinical signs”, “Radiography”,
90 “Respiratory Function Tests”, “Laryngoscopy”, “Ultrasonography”, and
91 “Bronchoscopy”. Various techniques have been studied to help clinicians assess
92 tube placement. However, despite 85 years of clinical practice, the search for higher
93 success rates and quicker intubation continues. Currently, chest radiography remains
94 the gold standard test to confirm tube position. However, rigorous evaluation of new
95 techniques is required to ensure the safety of newborn infants.

96
97

97 **Background**

98 Tracheal intubation remains a common procedure in the neonatal intensive care unit
99 (NICU) and the delivery room (DR)¹⁻⁵. The insertion of a tube into the trachea to
100 inflate the lungs of newborn infants has been routinely practiced for almost a century.
101 James Blundell described digital intubation in 1834 as “Inserting a tracheal catheter
102 guided along the fingers, through which the accoucheur could inflate the infant’s
103 lungs with his own breath”⁶. However, it took almost a century until Flagg described
104 his technique of introducing a metal tube into the trachea, using a small
105 laryngoscope incorporating an electric light⁷. Blaikley and Gibberd later suggested
106 using a rubber catheter instead of the rigid metal tube in 1935⁸. Until the 1970s,
107 tracheal intubation was considered a simple procedure once the anatomy and the
108 manipulation of the laryngoscope were understood^{9,10}. With the introduction of chest
109 radiography, tube position became clearly visible and malposition of the tracheal tube
110 was able to be detected¹¹.

111
112 The current gold standard for confirming tube position is a chest radiograph, however
113 this is often delayed until after ventilation via the tube has been commenced¹¹. Rapid
114 confirmation of correct tracheal tube placement at the point of care is important
115 because tube malposition is associated with serious adverse outcomes, including
116 hypoxemia, pneumothorax, lung collapse and death^{1-3,12-15}. This has led to the
117 development of new methods to confirm correct tube placement. Various techniques
118 have been studied to enable i) correct tube placement and ii) optimal position within
119 the trachea (Table 1). These techniques include i) clinical signs, ii) measurement of
120 tube length based on parameters of body size or gestational age, iii) fiber optic
121 techniques, iv) respiratory function, iv) chest radiograph, vi) capnography, vii)
122 ultrasonography, and viii) magnetic detection. However, rates of correct tube
123 placement, particularly for junior medical staff, are less than 50% and accidental
124 esophageal intubation is common^{1,2,5,16}.

125
126 The aim of this article was to review the available literature about tracheal tube
127 placement in neonates. Critical issues that must be addressed when correctly placing
128 a tracheal tube include i) correct placement of the tube in the trachea and not the
129 esophagus, ii) proper depth of the tube and iii) time to complete the procedure. The
130 article describes intubation in the NICU and DR. All the available techniques and
131 devices described can be used in both the NICU and DR during emergency and non-
132 emergency intubations.

133

134 **Search strategy**

135 We reviewed books, resuscitation manuals and articles from 1830 to the present with
136 the search terms “Infant, Newborn”, “Tracheal intubation”, “Resuscitation”, “Clinical
137 signs”, “Radiography”, “Respiratory Function Tests”, “Laryngoscopy”,
138 “Bronchoscopy”, and “Ultrasonography”. All languages were included. The full search
139 strategies for PubMed, EMBASE and PubMed Central are detailed in Appendix 1.

140

141 **Tracheal vs. esophageal tube placement**

142 Various techniques have been studied to determine correct tube placement including
143 i) clinical signs, ii) capnography, iii) respiratory function, iii) fiber optic techniques, and
144 iv) video laryngoscopy, which are described in detail below.

145

146 *Clinical signs*

147 International resuscitation guidelines recommend clinical signs and CO₂ detectors to
148 confirm correct tube placement¹⁷. Clinical signs of correct tube placement include a
149 prompt increase in heart rate, chest wall movement, visualization during direct
150 laryngoscopy of the tube passing through the vocal cords, presence of breath sounds
151 in the axillae and absence of breath sounds in the epigastrium and condensation in
152 the tube during expiration¹⁷. Direct laryngoscopy to observe passage of the tube
153 through the vocal cords was first described by Roberts in 1949⁹. Condensation of
154 water vapor in the tube lumen, although less likely with esophageal intubation, can
155 occur and hence is not a reliable sign¹⁸. In 1964, Doss reported his experience in an
156 essay on “Resuscitation of the Newborn”, stating that “The chest is auscultated while
157 blowing into the tube to verify that the latter is in the trachea”¹⁰. Bednarek et al., in
158 1975, reported 9/14 intubation of the right main bronchus: Four were without
159 complications, four caused atelectasis of varying degrees and one resulted in
160 bilateral pneumothoraces¹⁵. “Chest rise” is another clinical sign used to identify
161 correct tube placement. Doss described his observation of unilateral chest rise to
162 identify unilateral main bronchus intubation; “If only the right lung expands the tube is
163 withdrawn a little”¹⁰. More important, movement of the chest wall simulating
164 ventilation of the lungs can be seen with an esophageal tube¹⁸. Observational studies
165 in the delivery room during mask ventilation have shown that judgment of chest rise
166 is inaccurate^{21,22}. Several studies have reported that recognition of esophageal
167 intubation, using clinical signs alone, may take several minutes^{2,4,5,18,19}. In summary,
168 a number of clinical signs aim to identify correct tube placement but none have been
169 systematically studied. The best indicator that the tube is in the trachea remains the
170 clinical response when the patient shows a prompt increase in heart rate^{17,20}.

171

172 *Exhaled Carbon Dioxide (CO₂)*

173 CO₂ is exhaled from the lungs at concentrations much higher than present in air. It
174 can be detected using colorimetric devices or measured quantitatively using main-
175 stream, side-stream or micro-stream devices^{3-5,19,23-28}. Measuring exhaled CO₂ by
176 infrared gas analysis to monitor patients during anesthesia was first described in the
177 1960s^{23,24} (Table 1). However, using end-tidal CO₂ to detect correct tube placement
178 was first described in an animal model by Murray in 1983 and in human patients by
179 Linko in 1983^{25,26}. International resuscitation guidelines recommend clinical
180 assessment and exhaled CO₂ to confirm correct tube placement immediately after
181 intubation¹⁷. Although CO₂ devices are frequently used to assess tube placement²⁹⁻³²,
182 false negative results may occur, particularly when cardiac output is low³³ or when
183 the infant is in severe respiratory failure and the inflation pressure is not high enough
184 to ventilate the lungs^{5,34}.

185

186 Disposable semi-quantitative colorimetric CO₂-detectors are recommended in
187 international guidelines. With each inflation and expiration a pH-sensitive chemical
188 indicator undergoes color change (e.g. from purple to yellow), reflecting the change
189 in CO₂ concentration in the gas passing through it. Garey et al. reported the tidal
190 volume threshold achieve color change at the Pedi-Cap with >0.72 mL²⁷. An absence
191 of color change suggests that the tube is not in the trachea. However, false negative
192 findings may occur as listed above. Evidence comes from one observational and one
193 prospective cohort study (Table 2). Aziz et al. compared the Pedi-Cap[®] (Nelcor
194 Puritan Bennett, Pleasanton, CA) with clinical evaluation and radiography findings for
195 tracheal intubation in 45 newborn infants (24 (53%) in the DR and 21 (47%) in the
196 NICU) who required tracheal intubation. In that study, the Pedi-Cap[®] correctly
197 identified 30 tracheal tube placements and 12 esophageal tube placements (Table
198 2). However, in three cases a false negative result (no color change) was displayed⁴.
199 Schmölzer et al. compared flow signals with the Pedi-Cap[®] in 35 intubations (Table
200 2). The flow signal correctly identified all tube placements correctly, whereas the
201 Pedi-Cap[®] failed to change color on 12 occasions when the tube was correctly
202 placed⁵. Colorimetric CO₂-detectors may mislead clinicians intubating very preterm
203 infants in the delivery room and may fail to change color despite correct tube
204 placement in up to one third of cases⁵.

205

206 Systems with either main-stream, side-stream or micro-stream measurement of end-
207 tidal CO₂ have been described^{3,19,28} (Table 2). Repetto et al. compared end-tidal CO₂

208 with clinical assessment in the NICU and DR. The median times required for
209 capnographic and clinical determination of tracheal intubation were 9sec vs. 35sec,
210 and for esophageal intubation 9sec vs. 30sec¹⁹. Roberts et al. compared side-stream
211 capnography with clinical assessment in 100 intubations in 55 infants³. Capnography
212 rapidly and correctly identified esophageal tube placements in 39 of 40 infants (mean
213 1.6 sec). Capnography failed to identify successful tracheal intubation on only one
214 occasion. Clinical indicators of esophageal position were much slower (mean
215 97.1sec) and failed to identify successful tracheal intubation in 5 of 60 cases³. In
216 comparison, Hosono et al. demonstrated that compared to clinical assessment,
217 micro-stream CO₂-detectors correctly identified all tube placements, whereas clinical
218 assessment failed to identify three of eleven esophageal tube placements²⁸. We
219 need to emphasize that although micro-stream CO₂-detectors can be used to identify
220 correct tube placement, inexperience and lack of knowledge about the displayed
221 waveforms may lead to misinterpretation of the signals. Therefore, anyone using this
222 device must be trained to interpret capnography waveform signals (Figure 1). In
223 addition, all CO₂ devices may display false negative results particularly when cardiac
224 output is low or when the infant is in severe respiratory failure and the inflation
225 pressure is not sufficient to ventilate the lungs. Although observational studies have
226 evaluated side-stream or micro-stream CO₂-detectors, neither have been evaluated
227 in randomized trials in neonates.

228

229 *Respiratory Function Monitor*

230 In a Scottish medical negligence case from 1973, John Robinson first described the
231 use of airway pressure and gas flow waves to identify correct tube placement³⁵
232 (Table 1). Robinson reported that tidal volume measurements are feasible and
233 suggested that airway pressure and gas flow should be used to differentiate tracheal
234 from esophageal tube placement (Figure 1)³⁵. Almost any respiratory function
235 monitor can be used to measure and display airway pressure, gas flow and tidal
236 volume³⁶. A flow sensor placed between the tube and the ventilation device can be
237 used to determine tube placement³⁷. The operator should look for inspiratory and
238 expiratory gas flow waves. If gas flow can be observed in both directions, the tube is
239 in the trachea. If only inspiratory gas flow is observed, the tube is most likely in the
240 esophagus. Recently, Schmölzer et al. compared the gas flow signals to a
241 colorimetric CO₂-detector in an ovine model of neonatal resuscitation³⁷. Both
242 colorimetric CO₂-detector and flow signal identified all tube placements correctly.
243 However, the colorimetric CO₂-detector required at least three and up to 10 inflations
244 to identify tube location compared to one to two inflations with the flow sensor. The

245 available evidence during neonatal resuscitation derives from two observational
246 studies in the DR^{2,5} (Table 2). O'Donnell et al. compared clinical assessment (n=33
247 infants) and flow signal (n=7 infants) for identification of correct tube placement².
248 Clinical assessment of tube position took 39sec compared to 19sec using a flow
249 signal. Schmölzer et al. compared the gas flow sensor to a colorimetric CO₂-detector
250 during 35 intubation attempts of 20 infants during neonatal resuscitation⁵. In 21
251 (60%) intubations, both methods correctly identified successful tube placement, and
252 in 3 (9%) intubations both methods indicated the tube was not in the trachea.
253 However, in the remaining 11 (31%) intubations the colorimetric CO₂-detector failed
254 to change color despite the flow wave indicating correct placement. This study
255 suggests that colorimetric CO₂-detectors may fail to change color despite correct
256 tube placement in up to one third of cases and therefore may mislead clinicians
257 intubating very preterm infants in the DR. However, neither gas flow nor airway
258 pressure monitoring can distinguish tube placement in the trachea from one of the
259 main bronchi, which is a major limitation. Recently, in an animal model, electrical
260 impedance tomography was shown to correctly identify tube misplacement in the
261 bronchial tree³⁸. This device is currently best regarded as a research tool and is not
262 available to clinicians. Substantial training in the use of this technology is required to
263 ensure correct interpretation of the signals produced.

264

265 *Fiber Optic Devices*

266 A fiber optic laryngoscope/ bronchoscope for endotracheal intubation was first
267 described by Stiles in 1972³⁹ (Table 1). These devices emit light, and development of
268 the ultrathin bronchoscope has made flexible endoscopic intubation for the neonate
269 and small infant possible. Advantages include: i) easy identification of the carina or
270 the esophagus and ii) intubation can be performed either via nasal or oral route, high
271 correlation with chest radiograph³⁹⁻⁴². However, limitations and disadvantages include
272 i) desaturation during the procedure, ii) time required to visualize the vocal cords,
273 and iii) special training which is necessary to operate the flexible fiberscope
274 efficiently in emergency settings³⁹⁻⁴².

275 Fiber optic devices have been used to i) intubate newborn infants in the NICU⁴³ and
276 ii) assess correct tracheal tube placement^{40,41}. Finer et al. described non-emergency
277 endoscopic intubation of newborn infants in the NICU using a bronchoscope⁴³. The
278 majority of those infants were intubated prior to surgery or to investigate upper
279 airway abnormalities. There was evidence of transient hypoxemia during the
280 introduction of the bronchoscope into the trachea, but no infant had an oxygen
281 saturation lower than 85%⁴³. The longest part of the procedure was time required to

282 visualize the vocal cords adequately, and the most difficult aspect was the
283 introduction of the tip of the bronchoscope into the trachea⁴³. Although fiber optic
284 tracheal intubation is a reliable method of neonatal intubation and used in the NICU,
285 devices are not readily available in emergency situations. For this reason the
286 technique has not been adopted for delivery room intubations. This has led to the
287 development of fiber optic stylets, including viewing windows at the top of the shaft
288 stylet, miniature screens, and video laryngoscopes⁴⁴.

289

290 *Video Laryngoscopy*

291 Video laryngoscopy provides a view from the tip of the laryngoscope blade during
292 conventional direct laryngoscopy. These devices may use fiber optic image
293 transmission to a miniature screen or have an embedded small camera head.
294 Advantages of video laryngoscopes include: i) the image and light bundle does not
295 interfere with the operator's view of the airway, so the standard technique of
296 laryngoscopy remains possible, and ii) the light provided by the fiber optic bundle is
297 safe and does not cause burns⁴⁵⁻⁴⁹. Disadvantages include: i) increased weight of the
298 laryngoscope handle, and ii) failure of the light source, although this can occur with
299 any laryngoscope⁴⁵⁻⁴⁹. In the pediatric age group, video-assisted intubation was first
300 reported by Weiss in 2000 (Table 1)⁴⁵. In 100 children aged from 3 months to 14.3
301 years the trachea was successfully intubated on the first attempt without arterial
302 oxygen desaturation. Vanderhal et al. reported their preliminary experience using a
303 video laryngoscope⁴⁶. Although three intubations required more than three attempts
304 with the video laryngoscope, the overall number of intubation attempts was
305 reduced⁴⁶. Trevisanuto et al. reported video guided intubation using the GlideScope[®]
306 in five newborn infants⁴⁷. Intubation with the GlideScope[®] was successful in three of
307 the five cases, however in two cases direct laryngoscopy was performed after two
308 failed attempts with the GlideScope[®]⁴⁷. A randomized trial compared success rate of
309 correct tube placement for the GlideScope[®] video laryngoscope with direct
310 laryngoscopy⁴⁸ and reported no difference between the two devices⁴⁸. Although,
311 video laryngoscopes appear to be efficient and safe, randomized trials are required
312 to compare each video laryngoscope with a conventional direct laryngoscope, with
313 the other video laryngoscopes, and with the other types of intubation devices.

314

315 **Correct tube position within the Trachea**

316 Various techniques have been studied to determine optimal position within the
317 trachea including i) chest radiograph, ii) measurement of tube length based on

318 parameters of body size or gestational age, iii) external digital tracheal palpation, iv)
319 fiber optic techniques, and v) ultrasonography.

320

321 *Chest Radiograph*

322 A chest radiograph can be used to confirm correct tube position within the trachea,
323 which should be just below the level of the vocal cords and well above the carina.
324 Various techniques have been described to achieve tube positioning above the
325 carina prior to X-ray confirmation, but tube displacement, particularly into the right
326 main bronchus, remains common^{11,50}. While the current gold standard to assess
327 correct tube placement is chest radiography, no study has formally investigated the
328 use of chest radiographs to confirm tracheal tube placement in either the NICU or
329 DR¹¹. Guidelines for the interpretation of chest X-ray to assess correct tube
330 placement vary. Usually, the lower limit for the position of the tube is measured in
331 relation to the carina. The tip should be between 0.2 to 2 cm above the
332 carina^{11,13,15,51}. The medial border of the clavicles has been used in studies as
333 reference point, however the position of the clavicles can range from the 4th cervical
334 vertebrae to the 2nd thoracic vertebrae on chest radiographs⁵¹. In addition, studies
335 have shown that head extension or flexion from the neutral position can move a
336 tracheal tube by an average of 0.7 cm in either direction in newborns⁵¹. Hence using
337 the 1st thoracic vertebrae has been suggested to be an alternative reference point.

338

339 *Estimation of correct endotracheal tube insertion*

340 Several formulae have been proposed to achieve correct tube placement within the
341 trachea^{12,52-57}. In 1987 Yates et al. developed a formula based on tube lengths
342 recorded in their study (internal diameter x 3 + 2 cm)⁵². However, this formula has not
343 been tested in further studies. In 1979, Tochen described a simple formula to
344 determine the depth of tube insertion⁵³. The "tip-to-lip" distance or "7-8-9 rule",
345 estimates the depth (cm) of tube insertion to be $[1.17 * \text{infant's weight (in kg)} +$
346 $5.58]$ ⁵³. Using this formula an infant weighing 1kg would be intubated to a depth of
347 7cm, a 2kg infant to a depth of 8cm, and a 3kg infant to a depth of 9cm⁵³. However,
348 several studies have shown that the Tochen's 7-8-9 rule led to incorrect tube
349 placement in almost half of the infants studied. Peterson et al. studied the accuracy
350 of the 7-8-9 rule for tube placement in 75 consecutively intubated infants of less than
351 32 weeks gestation⁵⁴. Although correct tube placement was confirmed in infants >
352 750g body weight, tube depth was overestimated in infants < 750g⁵⁴. Amarilyo et al.
353 also assessed the reliability of the 7-8-9 rule during oro-tracheal intubation in 31
354 infants < 1000g¹². In almost half of the infants (47%), the rule resulted in incorrect

355 tube placement when checked by chest radiograph¹². Furthermore, Kempley et al.
356 recently re-examined Tochen's rule and reported that it assumes a linear relationship
357 between tube and body length and body weight^{56,57}. The relationship, although linear
358 for gestational age, is non-linear for body weight⁵⁷. Furthermore, the rule does not
359 apply for naso-tracheal intubation.

360

361 *External digital tracheal palpation*

362 The practice of tube palpation dates back to the time of its first use in 1834 by British
363 obstetrician, James Blundell, who used an tracheal tube to resuscitate an infant⁶.
364 Blundell noted that following digital intubation "the tracheal pipe can be felt on the
365 front of the neck whether the instrument is lying in the trachea or esophagus"⁶.
366 Bednarek et al. in 1975 reported that tracheal tube position could be confirmed using
367 digital suprasternal palpation in neonates and small infants¹⁵. Once the tube is
368 placed and the tip can be gently palpated in the suprasternal notch the position is
369 considered correct. Two studies compared external digital tracheal palpation to direct
370 laryngoscopy^{15,58}. Bednarek et al. reported that infants in the palpation group had
371 significantly more accurate tube placements, significantly lower rates of
372 complications, and no adverse effects compared to direct laryngoscopy¹⁵. In a
373 randomized controlled trial, Jain et al. showed that digital palpation correctly
374 predicted tube position on chest radiograph in 70% of included infants⁵⁸. Digital
375 palpation resulted in a small, non-significant improvement in rates of correct
376 placement compared with standard weight based criteria⁵⁸. The authors concluded
377 that suprasternal palpation appears to be a simple and teachable method of
378 confirming tracheal tube position in neonates when chest radiograph is unavailable,
379 and may be especially helpful during neonatal resuscitation prior to surfactant
380 administration.

381

382 *Fiber Optic Devices*

383 Fiber optic devices have been used to i) intubate newborn infants in the NICU⁴³ and
384 ii) assess correct tracheal tube placement.^{40,41} Vigneswaran et al. studied the use of
385 a 1.8mm fiberoptic bronchoscope for confirmation of tracheal tube position in 20
386 intubated infants⁴⁰. They concluded that the bronchoscopic technique of determining
387 tracheal tube position is as safe and accurate as the radiologic technique (correlation
388 0.91)⁴⁰. Shinwell et al. reported their experience in 65 infants⁴¹ and reported that the
389 accuracy of bronchoscopic measurement of tracheal tube tip position improved
390 markedly with user experience and correlated very well (0.96) with chest
391 radiograph⁴¹. Although fiber optic tracheal intubation is a reliable approach for

392 neonatal intubation and used in the NICU, devices are not readily available in
393 emergency situations.

394

395 *Ultrasound*

396 In 1986 Slovis et al. demonstrated that real-time ultrasound could be used to verify
397 correct tube position in newborn infants⁵⁹ (Table 1). Slovis et al. used the aortic arch
398 as a reference point and showed that tube position could be accurately determined in
399 18/21 (85%) intubations⁵⁹. Lingle et al. showed similar results using the same
400 technique⁶⁰. Galicinao et al. used two views with a linear transducer in infants and
401 children to verify correct tube placement⁶¹. Compared to physical examination +
402 colorimetric devices + chest radiography, bedside ultrasound was much faster⁶¹.
403 Dennington et al. (Table 2) reported good correlation between ultrasound and
404 radiograph measurements with most values being within 0.5 cm⁶². Advantages of
405 ultrasonography over radiography include: i) reduced radiation, ii) less handling,
406 particularly in critically ill infants, iii) potential to determine the tube position in the
407 delivery room particularly for early surfactant delivery and iv) earlier detection of
408 complications from malposition⁵⁹⁻⁶². Disadvantages of ultrasonography are i) the
409 specialized skills required, ii) difficulties to correctly identify anatomical landmarks
410 and iv) lack of widespread availability⁵⁹⁻⁶². These observational studies demonstrate
411 that ultrasound can be used to visualize tracheal tube position in infants. However, 1-
412 2 specially trained investigators performed all ultrasound evaluations in the reported
413 studies, which is a major limitation of these studies. And therefore it is not applicable
414 for daily clinical use. Randomized control trials are required before implementation of
415 this technique.

416

417 **Duration of the procedure**

418 Guidelines of the Neonatal Resuscitation Program developed by the American
419 Academy of Pediatrics recommend that the intubation procedure is completed within
420 30 seconds¹⁷. Two observational studies reported the times to intubate preterm
421 infants in the DR using a laryngoscope^{2,63}. Overall, 50-60% of intubation attempts
422 were successful, with the majority occurring within 30 seconds^{2,63}. Interestingly, two
423 studies reported that digital intubation was significantly faster compared to intubation
424 using a laryngoscope^{64,65}. However, digital intubation is not routinely used during
425 neonatal intubation.

426 Several studies have reported that experience significantly increased success rate of
427 tracheal intubation^{1,2,16,63}. O'Donnell et al. reported a significant difference between
428 grades of doctors, with more senior doctors intubating more rapidly². However, the

429 study by Lane et al. reported that fellows were on average 10 seconds faster than 1st
430 year residents or consultants⁶³. The main concern is the decrease exposure of
431 endotracheal intubation during residency and fellowship training^{1,16}. Leone et al.
432 reported that the mean number of intubation attempts per resident throughout
433 training decreased from 38 (\pm 19) in 1994 to 12 (\pm 6) in 2002¹. In addition, non-
434 invasive methods of respiratory support has been advocated which further reduces
435 the number of intubation attempts^{66,67}. Manikin studies reported significant
436 improvement in the time to visualize the vocal cords with the video laryngoscope
437 compared to direct laryngoscopy⁶⁸. However, success rate were similar within the
438 groups⁶⁸. In comparison, a randomized trial comparing intubation time for the
439 GlideScope[®] video laryngoscope with direct laryngoscopy reported no difference in
440 intubation time or success rate of intubation between the devices⁴⁸.

441

442 **Conclusions**

443 Currently chest radiography remains the accepted standard test to confirm tube
444 position. An increase in heart rate is the best indication of effective ventilation.
445 Exhaled CO₂ may give misleading, false negative results and readings should be
446 interpreted in conjunction with clinical signs. Respiratory function monitors, video
447 laryngoscopes, fiber optic devices and ultrasound to confirm tube position are
448 promising methods but require evidence from clinical trials before recommendations
449 can be made for their use in clinical practice.

450

451 **Figure legend:**

452 **Figure 1:** Figure 1A: Esophageal and tracheal flow waves to identify correct tube
453 placement comparison described by Robinson 1973³⁵. In comparison airway
454 pressure, gas flow, tidal volume and end-tidal CO₂ waves from a preterm infant born
455 at 29 weeks gestation requiring intubation in the delivery room. Figure 1B -
456 Esophageal intubation: Gas flow towards the infant is present. However, no
457 expiratory gas flow, tidal volume or end-tidal carbon CO₂ is displayed. Figure 1C –
458 Tracheal intubation: Gas flow towards and away from the infant is present. In
459 addition, tidal volume or end-tidal CO₂ are displayed.

460 Figure 1A: Reproduced from [Respiratory recording from the oesophagus,
461 Robinson³⁵], Br Med J. 1974; 4 (5938), 225, 2012] with permission from BMJ
462 Publishing Group Ltd.

463 Figure 1B & 1C: Reproduced from [Monitoring during neonatal transport, O'Reilly⁷¹,
464 Emergency Medicine 2012, epub] with permission from O'Reilly et al.

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761 **Timeline: Evolution of endotracheal tube placement in newborn infants**

- 1834 J. Blundell describes digital intubation and external digital tracheal palpation for correct tube placement for the treatment of Asphyxia Neonatorum⁶
- 1895 A. Kirstein invented the modern laryngoscope⁶⁹
- 1921 Sir I. Magill recommends the sniffing position for endotracheal intubation⁷⁰
- 1928 P.J. Flagg describes his technique of introducing a metal tube into the trachea using a small electrically lighted laryngoscope⁷
- 1935 J.B. Blaikley and G.F. Gibberd suggest to use a rubber catheter instead of the rigid tube⁸
- 1949 H. Roberts recommends "When intubation is required in an asphyxiated newborn infant, the endotracheal tube should be passed through the vocal cord by direct laryngoscopy"⁹
- 1964 A.D. Doss describes: "The chest is auscultated while blowing into the tube to verify that the latter is in the trachea"¹⁰
- 1964 Chest wall assessment to assess main bronchus intubation is described by A.D. Doss: "If only the right lung expands the tube is withdrawn a little"¹⁰
- 1966 Carbon dioxide monitoring during anesthesia is described by GW Burton²³
- 1971 L.R. Kuhns firstly describes tracheal tube position in infants using chest radiograph¹¹
- 1972 C. Stiles describes flexible fiber optic laryngoscope³⁹
- 1974 J. Robinson describes the assessment of tube placement using respiratory function³⁵
- 1979 - Calculations to correctly position an oropharyngeal and
onwards nasopharyngeal tube^{12,52-55,57}
- 1983 First report of capnography for detection of accidental esophageal intubation in adults by K. Linko²⁶
- 1986 First study describing Ultrasound to confirm correct tube position⁵⁹
- 1990 Magnetically detectable tracheal tubes¹⁴
- 1992 - Studies investigating colorimetric, main, side and micro-stream CO₂-
onwards detectors^{2-5,19,28}
- 2000 M. Weiss describes the video laryngoscopy technique for the intubation of newborns⁴⁵
- 2006 - Studies examining respiratory functions to detect correct tube
onwards placement^{2,5}
- 2007 - Studies examining ultrasound to detect correct tube placement^{61,62}
onwards

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764 **Table 2: Comparison of available studies**

Study	Design	Comparison	Number of intubations	Results	PP
External digital tracheal palpation					
Jain ⁵⁸	Randomized trial	Chest radiograph vs. suprasternal digital tracheal palpation	54	Digital palpation correctly predicted tube position by chest radiograph in 70%	75%
Exhaled Carbon dioxide					
Aziz ⁴	Observational	Clinical Assessment vs. PediCap [®]	45	CO ₂ correctly identified 30/33 tracheal, 12/12 esophageal tubes, 3/33 tubes wrongly identified by the PediCap [®] as in the esophagus, clinical assessment correctly identified all tubes (33 tracheal, 12 esophageal)	100%
Repetto ¹⁹	Observational	Clinical Assessment vs. Main-stream end-tidal CO ₂	27	CO ₂ and clinical assessment correctly identified 16/16 tracheal, 11/11 esophageal tubes	100%
Roberts ³	Observational	Clinical Assessment vs. Side-stream end-tidal CO ₂	100	CO ₂ correctly identified 59/60 tracheal, 40/40 esophageal intubations, and failed to identify 1 tracheal intubation; clinical assessment failed to identify 36/40 esophageal intubation	62%
Hosono ²⁸	Observational	Clinical Assessment vs. Micro-stream end-tidal CO ₂	54	CO ₂ correctly identified 40/40 tracheal, 14/14 esophageal tubes; clinical assessment failed to identify 3/14 esophageal intubation	93%
Respiratory Function Monitor					
Schmölzer ⁵	Observational	Flow sensor vs. PediCap [®]	35	Flow sensor correctly identified 32/32 tracheal, 3/3 esophageal tubes; PediCap [®] correctly identified 21/32 tracheal, 3/3 esophageal intubations; 11/32 tubes wrongly identified by the PediCap [®] as in the esophagus	100%
Ultrasound					
Lingle ⁶⁰	Observational	Ultrasound vs. Chest radiograph	9	Placement of 4 tubes correctly identified, 2 misplacement	100%
Dennington ⁶²	Observational	Ultrasound vs. Chest radiograph	29	Placement of 29 tubes correct identified	No ane

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Figure 1

