1 Introduction

2 Endotracheal intubation is commonly performed in horses undergoing general anaesthesia to provide oxygen, inhalation anaesthetic agents and other drugs. Moreover, 3 an adequate seal must be obtained to provide mechanical ventilation and avoid fluid 4 aspiration or environmental pollution with inhalant anaesthetics. Oedema and 5 haemorrhage of the laryngeal and tracheal mucosa, and tracheal stenosis and perforation 6 7 have been reported secondary to orotracheal intubation in horses (Heath et al. 1989, Touzot-Jourde et al. 2005, Saulez et al. 2013, Wylie et al. 2015). Consequences of these 8 9 may be fatal (Rainger et al. 2006, Wylie et al. 2015). In one study, 95% and 85% of 10 orotracheally intubated horses showed damage of the tracheal and laryngeal mucosa, 11 respectively; but these lesions were classified as mild and resolved without treatment 12 within 24 hours after extubation (Heath et al. 1989). This case report describes successful treatment of a horse with laryngeal and 13 14 tracheal trauma secondary to intubation during myelography and underscores the importance of monitoring carefully these horses for development of pneumonia. 15 16 **Case history** 17 A 10-year-old Warmblood mare weighing 608kg was admitted to the Royal Veterinary College hospital for investigation of poor performance and suspected intermittent 18 19 hindlimb ataxia. A neurologic examination performed at the time of admission 20 identified mild hypermetria in all 4 limbs but was otherwise unremarkable. No ataxia was observed. The orthopaedic exam was unremarkable. 21

22 Due to concerns over reports of intermittent ataxia and the implications for rider 23 safety, the owner elected for the mare to undergo head and cervical spine computed 24 tomography (CT) and myelography under general anaesthesia. The pre-anaesthetic 25 evaluation showed that the mare was bright and alert, in a good body condition score (6/9) with normal thoracic auscultation, respiratory rate, heart rate and rectal 26 temperature. Premedication consisted of acepromazine (Tranquinervin)¹ 0.02 mg/kg 27 intramuscularly and flunixin meglumine (Pyroflam)² 1.1 mg/kg intravenously one hour 28 and a half before the procedure. Then, the horse was premedicated with romifidine 29 $(\text{Sedivet})^3 0.08 \text{ mg/kg}$ intravenously. At this point, the mouth was rinsed with water 30

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² Norbrook Laboratories Limited, Northern, Ireland.

³ Boehringer Ingelheim Ltd, Bracknell, Berkshire, UK.

until clean. General anaesthesia was induced with ketamine (Ketamidor)⁴ 2.5 mg/kg and 31 midazolam (Dormazolam)⁵ 0.05 mg/kg intravenously. Once recumbent, the mare was 32 intubated with a lubricated 30 mm silicone tube and the cuff was inflated with a 30 ml 33 syringe until the indicator balloon felt firm to palpation. Then the horse was hoisted 34 onto the CT table, placed in left lateral recumbency, connected to a large animal 35 anaesthetic machine, and inhalational anaesthesia with isoflurane in oxygen and medical 36 air was initiated. At this point, a large persistent leak was identified despite attempts to 37 inflate the endotracheal tube cuff. The mare was extubated and reintubated with a 38 39 lubricated 26mm endotracheal tube. The initial endotracheal tube used showed a torn cuff. The mare was hoisted and repositioned several times until satisfactory position for 40 the CT scan was obtained meanwhile it was kept connected to the anaesthetic machine. 41 The horse was monitored with electrocardiogram, invasive blood pressure, end tidal 42 43 carbon dioxide, end tidal isoflurane concentration, inspired fraction of oxygen, arterial blood gas analysis and airway pressures. Arterial blood gas analysis revealed inadequate 44 45 partial pressure of Oxygen [PaO₂ 79 mmHg (FiO₂ 72%)] which was successfully corrected with a stepwise alveolar recruitment manoeuvre and 100 micrograms 46 47 salbutamol (Ventolin)⁶ inhaled via the Y-piece salbutamol port. Dobutamine⁷ was used to maintain mean arterial blood pressure above 60 mmHg. Following completion of the 48 CT scan and myelography, flexed and extended radiographs of the neck were obtained. 49 The mare was then hoisted into the recovery box, where the endotracheal tube cuff was 50 deflated. Once nystagmus appeared, romifidine (Sedivet)³ 0.02 mg/mg intravenously 51 was administered. The mare was extubated when swallowing reflex was present. 52 53 Recovery occurred uneventfully.

CT showed bilateral enlargement of the articular process joints of C4-C5, C5-C6 and C6-C7, with mild osteophyte formation and sclerosis of the subchondral bone. Mild enlargement of the articular process joints of C2-C3 and C3-C4 was also noted. There was no evidence of spinal cord compression. Figure 1 shows the position of the endotracheal tube cuff during CT scan.

59 Clinical findings

⁴ Richter Pharma AG, Wels, Austria.

⁵ Le Vet. Beheer B.V, Dordrecht, The Neederlands.

⁶ Glaxo Wellcome Production, Evreux, France.

⁷ Hameln pharma ltd, Gloucester, UK

- 60 Approximately 24 hours after general anaesthesia a persistent cough was noted, whilst
- 61 the vital parameters were unremarkable. Thoracic ultrasound identified an increased
- 62 number of comet tails and mild consolidation in both cranioventral lung fields
- 63 consistent with pneumonia. Upper airway endoscopy was performed, showing laryngitis
- 64 with moderate to marked inflammation and erosions extending caudally into the trachea
- 65 for approximately five cartilage rings. These lesions were symmetrical and localised at
- 66 8 o'clock to 10 o'clock and the 3 o'clock to 5 o'clock position.

67 Diagnosis

68 Laryngitis and moderative to severe erosive tracheitis with secondary pneumonia.

69 Treatment

- 70 The mare was treated with flunixin meglumine 1.1 mg/kg (Pyroflam)² intravenously
- every 24 hours, doxycycline⁸ 10 mg/kg orally every 12 hours for five days and a single
- dose of dexamethasone (Colvasone) 9 0.1 mg/kg intravenously. The following day,
- flunixin meglumine (Pyroflam)² was discontinued and phenylbutazone (Equipalazone)¹⁰
- 74 2.2mg/kg orally every 12 hours was started.

75 Outcome

- 76 The mare remained bright and normothermic over the following 72 hours. Upper airway
- endoscopy and thoracic ultrasonography showed a marked improvement of the
- 78 laryngeal and tracheal inflammation with resolution of the previously identified
- consolidation in the cranioventral lung fields. The mare was discharged from the
- 80 hospital four days after general anaesthesia, at which time the cough had resolved.

81 Discussion

- 82 The most common cause of pneumonia in adult horses is aspiration of microorganisms
- 83 inhabiting the nasopharynx or oral cavity. However opportunistic bacteria can colonize
- the lungs when pulmonary defence mechanisms are compromised, by, for example,
- 85 general anaesthesia (Reuss and Giguere, 2015). Moreover, intubation results in
- 86 enhanced capacity of gram-negative bacteria to bind tracheobronchial cells, favouring
- colonization and pneumonia. Injured airway creates binding sites for bacteria in the

⁸ Bova Compounding, Caringbah, Australia.

⁹ Norbrook Laboratories Ltd, Corby, UK.

¹⁰ Dechra Limited, Skipton, UK

bronchial tree and mucus production creates potential sites for bacterial adherence
(Levine 1991, Raidal 1995, Rainger et al. 2006).

Laryngeal trauma is common after orotracheal intubation. In one study laryngeal 90 trauma occurred in 35 out of 38 intubated horses (Heath et al. 1989). Of those horses, 91 92% had mild to moderate focal mucosal ecchymosis but these lesions improved 24 92 93 hours after extubation without intervention. The case presented by the authors had to be reintubated because a persistent leak was identified. Likely, the endotracheal tube cuff 94 95 was ripped by the teeth during intubation. Although no studies have evaluated the 96 consequences of reintubation, the degree of difficulty placing the tube has not been 97 associated with increased laryngeal and tracheal damage (Heath et al. 1989). Therefore, 98 we considered less likely that re-intubation was the cause of the extensive lesions found in the laryngeal and tracheal mucosa. 99

100 Tracheal damage after intubation is reported in horses (Touzot-Jourde et al. 2005, Saulez et al. 2013, Wylie 2015, Miller and Auckburally 2018). Tracheal necrosis 101 102 is related to a reduction of the capillary blood flow in the tracheal mucosa and has been 103 associated with fatality due to secondary tracheal stenosis and infection (Wylie et al. 104 2015) and pleuropneumonia (Rainger et al. 2006). There are several causes for tracheal 105 damage including use of large endotracheal tubes, over-inflation of the endotracheal cuff, positioning of the endotracheal cuff near the thoracic inlet, changes in the position 106 of the horse's neck with the endotracheal cuff inflated and chemically induced injuries 107 (Burns 2019). In this case report, movement of the endotracheal tube was considered as 108 109 the most likely cause of the tracheal damage due to the extension, location and 110 symmetry of the lesions. However, cuff inflation or overinflation could have also played a role. In humans, movement of the endotracheal tube during patient head and neck 111 112 positioning is described (Yap et al. 1994, Tailleur et al. 2016). In one study, a 30 113 degrees flexion from neutral position moved the tip of the endotracheal tube an average of 5.5 mm closer to the carina, while a 30 degrees extension moved the endotracheal 114 115 tube an average of 6.3 mm away from the carina (Yap et al 1994). One study in dogs found that flexion and hyperextension of the neck leads to movement of the 116 117 endotracheal tube within the trachea up to four vertebral spaces (Quandt et al. 1993). 118 Moreover, extension of the neck causes an increase of length of the trachea and a 119 decrease of the cross-sectional axis, increasing the appositional forces between the tracheal mucosa and cuff wall (Burns 2019). Additionally, in dogs, movement of the 120 121 endotracheal tube can also lead to endobronchial intubation, and laryngeal and tracheal

damage due to the shearing forces between the tracheal mucosa and cuff wall (Quandt et 122 123 al. 1993). Movement of the endotracheal tube in the adult horse is unlikely to cause 124 endobronchial intubation due to relative length of endotracheal tube compared to 125 tracheal length. One case report described tracheal lesions and pleuropneumonia leading to fatality after myelography in a horse (Rainger et al. 2006). The authors also 126 127 postulated that the extension and flexion of the neck during myelography were the cause of the tracheal damage. Differences exist with our case report as our diagnosis was 128 performed antemortem based on diagnostic findings and clinical signs, location and 129 130 distribution of the tracheal damage and clinical outcome. The use of endotracheal tubes has been linked to tracheal damage when cuffs are inflated until a seal is obtained 131 132 during mechanical ventilation or even when low cuff inflation pressures have been used (Heath et al. 1989, Touzot-Jourde et al. 2005). Therefore, we cannot conclude if cuff 133 134 movement or cuff inflation, or a combination of both, were the main cause of the tracheal damage. 135

136 One study evaluated the effect of two different endotracheal tubes on tracheal trauma (Heath et al. 1989). The majority of the horses (95%) intubated with a cuffed 137 138 endotracheal tube showed tracheal lesions. These lesions were mainly from 1 to 1.5 cm² in the ventral tracheal wall, while lesions in dorsal and lateral tracheal wall were rarely 139 reported (Heath et al. 1989). Larger lesions, 10 to 15 cm, were found at the level of the 140 endotracheal tube cuff. The use of cuffless endotracheal tubes, was also associated to 141 the presence of tracheal (83%) and laryngeal (100%) damage. In the case presented 142 143 here, lesions were found caudal to the larynx in a symmetric shape, being compatible with the endotracheal tube cuff position and shape. 144

145 In the author's institution silicone tubes are used. These endotracheal tube cuffs 146 are classified as low volume-high pressure. In this type of cuffs measurement of the 147 endotracheal tube cuff pressure does not equal the tracheal wall pressure, and devices used to measure intracuff pressure are not useful to estimate the tracheal wall pressure 148 149 (Muir and Hubbell 2009). One study that used similar silicone tubes than in this case report evaluated the relationship between tracheal damage and intra-cuff pressures, 150 151 reported mild epithelial attenuation and erosion as the most common lesion in the trachea of horses in which the endotracheal tube cuff was inflated to seal a leak with 152 153 cuff pressures varying between 80-100 cm H₂O, and more severe damage with intracuff pressures of 120 cm H₂0 (Touzot-Jourde et al. 2005). Therefore, we consider 154 155 unlikely that cuff over-inflation caused the extensive and severe tracheal damage, as the cuff was inflated, in the second intubation, until a leak was sealed to allow mechanical
ventilation. Moreover, both endotracheal tubes were checked prior to induction of
general anaesthesia and no abnormalities were detected in the cuff inflation or shape.

As routine protocol in our institution, the endotracheal tube is cuffed after intubation until the pilot balloon feels firm. The cuff is further inflated if a leak is detected during mechanical ventilation. Once general anaesthesia is finished, horses are hoisted to the recovery box with the cuff inflated to avoid aspiration of gastrointestinal content. Once in the recovery box, the cuff is deflated. It has been shown that pilot balloon palpation is an inaccurate way of assessing the cuff inflation in dogs, and monitoring cuff pressures with a manometer is recommended (Briganti et al. 2012).

In order to avoid the tracheal damage induced by movement of the cuffed endotracheal tube, the animal could have been disconnected from the anaesthetic machine and the cuff deflated for hoisting or neck flexion and extension. During this period of time, total intravenous anaesthesia could have been used to maintain general anaesthesia.

171 Pneumonia was diagnosed based on clinical signs and thoracic ultrasonography. 172 Whilst tracheal wash and culture is recommended, it was not performed in this case as 173 the horse improved following empirical administration of doxycycline. Doxycycline was selected as it has good penetration and concentration in lung tissue and has broad 174 spectrum activity against respiratory pathogens (Womble et al., 2007). Anti-175 inflammatory drugs are also recommended for the treatment of pneumonia (Wilkins and 176 Lascola, 2015). In this case, phenylbutazone was administered due to its anti-177 178 inflammatory, analgesic and antipyretic properties. A single dose of dexamethasone was 179 given to suppress tracheal inflammation and ameliorate the cough.

180

181 Conclusions:

Orotracheal intubation is recommended and commonly performed in horses. Tracheal
and laryngeal trauma is not uncommon and this may lead to pneumonia. Horses should
be monitored closely for signs of tracheal and laryngeal trauma after general anaesthesia
for myelography.

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