

Vocal cord palsy as a sequela of paediatric cardiac surgery – a review

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Abstract:	<p>Background: Vocal cord palsy (VCP) is a recognised complication of cardiac surgery in the paediatric population and can lead to debilitating complications. Whilst there is an abundance of literature highlighting the presence of this complication, there is a scarcity of research focusing on the pathophysiology, presentation, diagnosis, and treatment options available for children affected by VCP.</p> <p>Materials and methods: Electronic searches were conducted using the search terms: "Vocal Cord Palsy", "VCP", "Vocal Cord Injury", "Paediatric Heart Surgery", "Congenital Heart Surgery", "Pediatric Heart Surgery", "Vocal Fold Movement Impairment", "VFMI", "Vocal Fold Palsy", "PDA Ligation". The inclusion criteria were any articles discussing the outcomes of VCP following paediatric cardiac surgery.</p> <p>Results: The available literature surrounding the topic have been summarized within their appropriate sections. The two main populations affected by VCP are paediatric patients undergoing aortic arch surgery or those undergoing PDA ligation. There is paucity of prospective follow-up studies; it is therefore difficult to reliably assess the current approaches and the long-term implications of management options.</p> <p>Conclusion: VCP can be a devastating complication following cardiac surgery, which if left untreated, could potentially result in debilitation of quality of life and in severe circumstances could even lead to death. Currently, there is not enough high-quality evidence in the literature to</p>

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	aid recognition, diagnosis, and management leaving clinicians to extrapolate evidence from adult studies to make clinical judgements. Future research with a focus on the paediatric perspective is necessary in providing evidence for good standards of care.

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Vocal cord palsy as a sequela of paediatric cardiac surgery – a review

Running title: Nerve injury

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Abstract:

Background: Vocal cord palsy (VCP) is one of the recognised complications of complex cardiac surgery in the paediatric population. Whilst there is an abundance of literature highlighting the presence of this complication, there is a scarcity of research focusing on the pathophysiology, presentation, diagnosis, and treatment options available for children affected by VCP.

Materials and methods: Electronic searches were conducted using the search terms: "Vocal Cord Palsy", "VCP", "Vocal Cord Injury", "Paediatric Heart Surgery", "Congenital Heart Surgery", "Paediatric Heart Surgery", "Vocal Fold Movement Impairment", "VFMI", "Vocal Fold Palsy", "PDA Ligation". The inclusion criteria were any articles discussing the outcomes of VCP following paediatric cardiac surgery.

Results: The two main populations affected by VCP are paediatric patients undergoing aortic arch surgery or those undergoing PDA ligation. There is paucity of prospective follow-up studies; it is therefore difficult to reliably assess the current approaches and the long-term implications of management options.

Conclusion: VCP can be a devastating complication following cardiac surgery, which if left untreated, could potentially result in debilitation of quality of life and in severe circumstances could even lead to death. Currently, there is not enough high-quality evidence in the literature to aid recognition, diagnosis, and management leaving clinicians to extrapolate evidence from adult studies to make clinical judgements. Future research with a focus on the paediatric perspective is necessary in providing evidence for good standards of care.

1. Introduction:

Congenital heart disease afflicts approximately 0.8-1.2% of live births worldwide [1-3]. A third of the defects can be expected to be severe and present as candidates for cardiac surgery [4,5]. Vocal cord palsy (VCP) is a recognised and important complication in children which can follow congenital cardiac surgery. VCP may result when either one or both of the recurrent laryngeal nerves (RLNs) are damaged; these children may present with speech, feeding, and respiratory impediments which are exaggerated in bilateral VCP (BVCP) [6].

Iatrogenic injury to the nerve fibres during an invasive surgical intervention may result in the paralysis of unilateral or bilateral vocal cords [7]. Daya et al. described that left-sided VCP, was more common than bilateral VCP following paediatric cardiac surgery [6].

The risk of VCP occurring is contingent on patient and surgical aspects. Patient-related factors include female gender, prematurity, and low birth weight [8,9]. Additionally, aortic arch manipulations and PDA ligations pose the most significant procedural risks of VCP [10]. Anticipating these factors better prepares the clinician for multidisciplinary collaboration - especially with otolaryngology and anaesthesiology for the management of emergency airway obstruction.

It is particularly important to consider the variable complications, long-term outcomes, and quality of life for the VCP patient. There will invariably be differences in the extent to which the nerves and vocal cords are affected between patients, and by extension, the degree to which phonatory and respiratory functions are affected. In an acute setting, airway obstruction is a life-threatening complication of VCP and if not treated can lead to aspiration, respiratory distress, and even death [10]. In the longer-term, a developing child's ability to engage in conversation or eat without the need of feeding tubes can all be compromised. This can also be detrimental to both the child and their family's quality of life [11]. Appropriate knowledge on VCP is important as it can guide the diagnosis and management. Additionally, many areas of the multidisciplinary team such as cardiology, otolaryngology, anaesthesia, and speech-language therapy can benefit from this high-quality evidence. Additionally, there are very few long-term follow-up studies that have assessed the quality of life and long-term impacts of children affected, making it difficult to dictate necessary clinical interventions following discharge.

The aim of this review is to summarise and evaluate the available evidence surrounding VCP following cardiac surgery in the paediatric population, with a broader focus on management strategies and the impact on quality of life for these patients.

2. Clinical assessment of patients with VCP:

The clinical presentation of a child with VCP can be split broadly into features affecting phonation and features affecting respiration. The most common, and sometimes the only feature of VCP, is the presence of stridor. Bilateral vocal cord palsy (BVCP) has been reported to have a higher incidence and more severe presentation of stridor compared to unilateral VCP (UVCP). Other symptoms include a weak cry, dysphagia, and potential

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3 aspiration. Intercurrent cardiac and neurological disorders can also complicate the clinical
4 picture and cause or contribute to respiratory distress [6].
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6 While the occurrence of VCP following cardiac surgery is rare, it is important to note that
7 this is heavily reliant on the child's clinical examination. As many patients may be
8 asymptomatic or only present with subtle clinical signs after a certain period of time, they
9 may have not been adequately investigated initially [10]. Investigations are either invasive
10 or non-invasive, and each carry associated advantages and disadvantages to both the child
11 and clinician.
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14 2.1: Non-invasive methods

15 A non-invasive method that can be undertaken at the bedside is the use of ultrasound to
16 study vocal cord mobility. Shaath et al [12] conducted a study with 10 children who had
17 undergone heart surgery and used ultrasound to assess the movements of the vocal cords.
18 They found it was successful in recognising when VCP was not present. However, it was not
19 as successful in diagnosing VCP as there could be some movement of the vocal cords despite
20 being paralysed [12]. Ultrasound is normally used as an adjunct to endoscopy rather than as
21 an alternative.
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26 2.2: Invasive methods

27 Laryngoscopy is an invasive intervention that is used for vocal cord visualization. Literature
28 suggests that fiberoptic flexible laryngoscopy (FL) should be utilised when investigating VCP
29 and may be the only diagnostic option necessary as it allows clear visualisation of the vocal
30 cord dynamics and the patency of the airway [13].
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33 Direct rigid laryngoscopy can be conducted under general anaesthesia. A competitive
34 advantage over FL is that the arytenoids can be palpated to exclude cricoarytenoid joint
35 fixation, which is another differential of VCP [6]. However, the palpation should be done
36 cautiously to avoid iatrogenic injury which could lead to prolonged intubation. The use of
37 either FL or rigid laryngoscopy should be carried out regularly to assess the status of the
38 vocal cords and to monitor after treatment for improvement [14].
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42 Laryngeal electromyography (LEMG) can also be utilised to assess recurrent laryngeal nerve
43 function. Monitoring electrodes are attached to the thyroarytenoid muscles intraoperatively
44 to look at the extent of immobility. A retrospective study by Scott et al. [15] found that in
45 50% of patients, the results from LEMG resulted in a change in management for the patient.
46 Additionally, Maturo et al. [16] built on Scott et al.'s [15] work and used LEMG as a method
47 to predict if recurrent laryngeal nerve function would return. In children who had
48 undergone PDA ligation, they found that the absence of action potentials by 6 months
49 meant it was unlikely that the VCP would resolve [16].
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53 Having expressed that routine diagnosis ought to replace symptomatic diagnosis, the utility
54 of diagnostic methods must be considered. Whilst FL is the preferred choice, it can be
55 challenging to perform in young children. Nonetheless, it performs better than direct
56 laryngoscopy - which requires a greater level of user expertise and in smaller neonates, it
57 may require general anaesthesia. Non-invasive methods such as ultrasound are also very
58 user-dependent and may not be opted for in severe or life-threatening presentations of
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VCP. The diagnostic method is crucial in the assessment and treatment of VCP; not only does it dictate the next step but can also be used to predict the outcome for the child and hence the quality of life. Regular use of FL is recommended to assess the patency of the airway and vocal cord status.

3. Management:

Management options for VCP can be split into two options: medical and surgical. Following diagnosis, the clinician will be able to evaluate the need, type, and urgency of treatment.

3.1: Medical Management

Before deciding the modality of treatment, the child can be observed through clinical observation and serially measuring oxygen saturations to check for respiratory distress [6]. If the child shows signs of airway compromise, more urgent intervention is required. Spontaneous recovery is an option for VCP in stable children where a 'wait and watch' method is adopted. The mechanism of this process is not well understood however, in UVCP it is hypothesised that the healthy vocal cord can compensate for the other until neurological regeneration occurs [17]. During this time, the use of a nasogastric tube is common until the child is able to adequately cough and swallow [18].

Another option is speech therapy which has been found to be successful, particularly in older children. While there is a lack of reported evidence for VCP following cardiac surgery, 84% of patients with neurological malignancies found improvement in both speech and swallowing function. Further clinical trials are necessary to ascertain if this remains true for children with VCP following cardiac surgery.

Steroid therapy is favoured due to its likelihood of helping to reduce post-operative voice changes by preventing surgery-related complications such as oedema, which are linked to voice change [19]. Aside from their analgesic, prophylactic and antiemetic properties, steroids are also able to promote post-paresis nerve function recovery [20] and are used as part of the treatment of hoarseness [21]. A single-dose of intraoperative steroids can reduce the duration of temporary RLN palsy; however, it does not reduce the rate of RLN palsy [20] nor does it have a beneficial effect on voice-related quality of life [19].

3.2: Surgical Management

Surgical management is undertaken after the recommended 8-12 months period of observation; unless the child is initially unstable and requires early surgical intervention to protect the airway. Findings from Jabbour et al. [22] and Misono et al. [20] indicate that between 20-40% of children remain symptomatic with UVCP and emerge as candidates for surgical intervention. The main surgical interventions for UVCP are medialisation (injection laryngoplasty or type 1 thyroplasty) and non-selective recurrent laryngeal innervation and as shown in Table 1, each procedure carries its own set of advantages and disadvantages [21]. The most popular BVCP interventions are tracheostomy and iterations of posterior cordotomy [23].

3.2.1: Surgical management for UVCP

Injection Laryngoplasty

This involves injecting a synthetic material into the paralysed cord to medialise it [21]. The aim is to allow phonation and prevent aspiration, as a temporizing intervention either to mitigate symptoms or in anticipation of spontaneous reinnervation [24]; it is also used in the immediate postoperative period prior to non-selective laryngeal nerve innervation [21]. It has great utility in treatment plans aiming to initiate and/or advance an oral diet in paediatric patients with surgically induced VCP, especially if conducted within 6 months of the surgical injury [25]. There are a variety of materials available with variable immunogenicity such as Teflon, hyaluronic acid, collagen [21], and carboxymethylcellulose, which despite lasting between 1-2 months, is a strong material option as it has a safe biological profile with few reported complications and good outcomes in voice improvement [21].

Type 1 Thyroplasty

Type 1 Thyroplasty is a technically demanding and invasive procedure involving the insertion of synthetic implants through a window fashioned in the thyroid cartilage. The materials used are either Silastic, cartilage, or Gore-Tex [20, 26], for which there are less reported immunogenic effects than for injection laryngoplasty materials [16]. Butskiy et al. [27] reported 88% (n=8) improvement or recovery from aspiration in children and this finding could be the substrate of further clinical study amongst larger cohorts. Revisions are necessary, however, in paediatric patients to accommodate the growing and developing laryngeal anatomy [21]. The anticipated amount and frequency of these revisions is difficult to deduce from the scarcity of available long-term follow-up data [27]. In any case, delayed access is understood to cause increasingly greater technical challenges, especially if conducted post-puberty [28]. Owing to its failure to correct persistent phonatory gaps [21], Isshiki et al. [29] first described arytenoid adduction as a relatively simple yet critical adjunct to Type 1 Thyroplasty, despite its leading to longer procedural times and being technically challenging [30].

Reinnervation

Non-selective recurrent laryngeal reinnervation (NSLR) of the abductor and adductor intrinsic muscles of the larynx is achieved by anastomosing the RLN to itself or an adjacent motor nerve [23]. In comparison to static medialisation thyroplasty, NSLR confers a higher voice quality due to better vibrational capacity of the vocal cords [31, 32] with more long-term stability owing to preserved laryngeal muscle bulk and tone [31]. It confers a significantly reduced risk of aspiration in children due to the improved glottic closure patterns [24]. It has been recommended that the optimal time for NSLR is after 3 years of age [21]. Although general anaesthesia [33] obviates the need for intraoperative patient cooperativity, there are concerns related to the neurocognitive harm caused by prolonged anaesthetic time [34].

The independence from synthetic materials allows laryngeal development and removes the necessity for revision surgeries that are associated with thyroplasty. The procedure is understood to be safe without a significant risk of airway or wound complications [33, 35], but a consultation with a paediatric cardiologist would be useful in any case when assessing operative risk. Unlike the medialisation techniques, the improvement seen with UVCP is not immediate. Findings from Smith et al. [33] – which shows consistency with other studies

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3 from a systematic review of laryngeal reinnervation techniques [35] - suggest that a positive
4 improvement from reinnervation can be expected after a minimum of 6 months with stable
5 results by 12 months. Having already undergone previous cardiac surgery, patients (and/or
6 their parents) may show resistance to the prospect of further surgery due to additional
7 scarring [24] and so ought to be counselled appropriately on the implications and long-term
8 benefits. Ongkasuwan et al [36] suggests that NSLR procedures should be performed as
9 early as possible to optimise outcomes but also noted that favourable outcomes could still
10 be achieved two decades after the onset of childhood neuronal VCP. They also speculate
11 that preoperative LEMG may have utility in predicting voice outcomes following NSLR but
12 that further data is required [36].
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17 3.2.2: Surgical management for BVCP

18 *Iterations of posterior cordotomy*

19 This is an irreversible procedure that improves the laryngeal airway by removing a posterior
20 section of the vocal fold, while potentially sacrificing voice and/or swallowing ability [23].
21 Despite modification by otolaryngologists in recent years to mitigate these compromises,
22 the treatment approach remains suboptimal [37].
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26 *Tracheostomy*

27 A tracheostomy involves either a temporary or permanent opening of the trachea. This gives
28 relief of airway obstruction and protection from aspiration but does not improve phonation
29 (and will remove any ability to phonate unless fenestrated) [38]. Despite a tracheostomy
30 improving the airway [23], findings by Westwood et al. [39] substantiate that there is an
31 inherent compromise on quality of life for both the patient and their caregivers and
32 recommend psychosocial support for families. The main problem is the stoma that requires
33 continual care [40] and so research of less-invasive techniques is ongoing [38].
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37 *Selective reinnervation*

38 In contrast to NSLRI, selective recurrent laryngeal innervation of the posterior
39 cricoarytenoid muscles - involves specifically reinnervating distal abductor branches of the
40 RLN [23]. Crumley [41] was instrumental in defining the phrenic nerve as a donor target for
41 the abductor muscle bellies. A case series by Lee et al. [42] enforced a belief of the
42 technique's safety and appropriate use in children as young as 2 years old. Evidence remains
43 largely limited to adult patients however and further studies with larger sample sizes are
44 warranted for this promising technique.
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48 *Arytenoidectomy*

49 This is the irreversible surgical removal of the arytenoid cartilage to enlarge the airway [30].
50 Advances in arytenoidectomy procedures started with the use of endoscopic lasers, which
51 resulted in increased precision and decreased oedema post-surgery, and most importantly,
52 did not require tracheostomy to be performed [38]. However, surrounding tissues can be
53 damaged by the heat from the lasers, leading to increased susceptibility to granuloma and
54 scar formation, and thus resulting in more revision surgeries as the airways will inevitably
55 re-narrow [44]. Arytenoidectomy can be combined with arytenoid cordectomy or used
56 alone, yet both procedures can adversely affect voice quality [37].
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Laterofixation (Suture lateralisation/laterofixation)

Laterofixation of the vocal cord and/or arytenoid cartilage is indicated in BVCP patients as an alternative to tracheostomy or as a temporising measure when the recovery of laryngeal function is expected [44]. It reversibly enlarges the airway without damaging phonatory tissues so that voice can be later be restored and it can be performed independently or concurrently to endoscopic procedures [37]. Good success rates have been reported in paediatric cohorts [44] with superior performances in aerodynamic measures when compared to irreversible static procedures (e.g., cordotomy and arytenoidectomy); nonetheless, complications including aspiration, hoarseness, and need for adjustments should be anticipated [37].

Laryngeal pacing (Functional electrical stimulation)

Laryngeal pacing is an upcoming surgical option that has the potential to confer high-quality ventilatory improvement in the absence of swallowing and phonatory compromise [37]. As demonstrated by Mueller [45], the laryngeal pacing system involves electrode stimulation of the posterior cricoarytenoid (PCA) muscle(s) with the implant being situated in a subcutaneous pocket of the sternum. Candidates with aberrant RLN reinnervation will optimise the utility of this procedure [45]. Future research ought to address the absence of paediatric data for this technique [38]. The actual procedure itself is complicated and also proves expensive as the device needs replacement at least every 10 years [46].

Botulinum injection

The toxin botulinum is also used in the treatment of BVCP, especially in patients with laryngeal synkinesis, which is the involuntary movement of muscles following voluntary movement in one muscle. It promotes ventilation by causing flaccid paralysis in adductor muscles through inhibiting acetylcholine from being released from axon terminals [37]. Once adductor inspiratory motor neurons and muscles are blocked, abductor inspiratory motor neurons are able to produce a glottal opening, thereby enabling ventilation. Unlike cordotomy and arytenoidectomy, this treatment option has little to no effect on the voice. However, it is only a temporary fix as repeated injections are required every 3 months [37].

3.3: Future management options:

Further research could offer enhanced detail into our understanding of VCP and allow better diagnosis and treatment. Alternative therapies that promise regeneration of the laryngeal nerves and muscle trophism is gene therapy [37]. Gene therapy offers exciting prospects; whilst untested in the paediatric community, gene transduction is in pre-clinical trials for adult patients. Vectors can be delivered to certain points of the body such as the nervous system, musculature, or mucosa around the larynx. This could provide protection to the neurons, regeneration of the axons, and also prevention of muscle degeneration [47]. However, while gene therapy is under consideration for treatment of BVCP caused by neurodegenerative conditions, it does not prevent synkinesis [37]. Gene therapies also pose a risk of neuronal damage via the delivery of viral vectors into the CNS [38]. Gene therapy is still in the experimental stages, and more research is needed to test the effectiveness of

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3 neuronal regeneration and to find solutions to their adverse effects, before it can be
4 considered as treatments for iatrogenic BVCP, especially in the paediatric population.
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7 **4. Outcomes:**

8 The outcome of patients with VCP post-cardiac surgery varies with respect to a number of
9 factors such as site (either unilateral or bilateral), gestational age, and surgical procedure.
10 The complications following VCP can either be transient or permanent and vary widely in
11 severity [14].
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14 4.1 Recovery:

15 A recent systematic review by Engeseth et al. [48] reported low left VCP recovery rates (0-
16 33%) from a range of PDA ligation studies (17 retrospective; 4 prospective) which exposes
17 infants to life-long complications. Table 2 supports that the recovery of vocal fold function
18 following congenital cardiac surgery is subject to substantial variation based on surgical
19 procedure and gestational age. The highest rate of recovery is seen with aortic arch repair,
20 specifically the Non-Norwood procedures with 86%, while the lowest rate of recovery is
21 seen in PDA ligation with 0% [49, 50]. Supportive care should be implemented during the
22 postoperative period to aid the recovery of VCP [10]. It is also important to observe that a
23 variation exists in the length of follow-up periods for VCP recovery across different studies
24 which range between 3 – 16.4 months in Table 2. Many studies have relatively short-follow
25 up periods which may prompt underestimated VCP recovery rates - there could be under
26 detected greater rates of resolution within asymptomatic cohorts [50]. Additionally, without
27 conducting pre- and postoperative FL, there are inconsistencies in the evaluation of vocal
28 cord motion before and after therapeutic intervention which challenges the comparability
29 of studies.
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35 Truong et al. [50] observed that premature infants, especially neonates (≤ 28 weeks
36 gestational age), had lower rates of recovery of vocal cord function in comparison to infants
37 born at term across a variety of cardiac surgeries. Evidence from Zbar et. al [52] suggests
38 that infants of greater prematurity and lower birth weight are more likely to develop left
39 VCP during PDA ligations. Birth weight also appears to play a focal role in the incidence of
40 left sided VCP following PDA ligation; a greater incidence was found in infants with a lower
41 birth weight (5 of 22; 22.7%), compared to the incidence for heavier babies (1 of 46; 2.2%)
42 [52]. Infants that weigh ≤ 1 kg at birth are defined as extremely low birth weight (ELBW) and
43 it is preferential to use clips during PDA ligation in this population to lessen the extent of
44 dissection and mitigate the risk of injury to the great vessels. However, using clips is
45 associated with higher rates of VCP; it can result in RLN injury due to the close proximity of
46 the RLN to the ligation site [52]. Meticulous monitoring of ELBW infants undergoing PDA
47 ligation is imperative - Benjamin et al. [14] found an increased likelihood of
48 bronchopulmonary dysplasia, reactive airway disease or insertion of a gastrostomy tube in
49 this cohort.
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55 4.2 Adverse Outcomes

56 Engeseth et. al [48] highlighted that left VCP is associated with a range of adverse outcomes
57 among infants. There is a wide range of adverse outcomes spanning respiratory outcomes
58 and comorbidities such as bronchopulmonary dysplasia, laryngeal outcomes (dysphonia,
59 laryngomalacia, subglottic stenosis), feeding outcomes (gastrostomy, tube feeding) and
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3 other outcomes (hospital stay, readmission rate) [48]. For example, a retrospective study by
4 Nichols et al. [53] reviewed the functional outcomes with vocal fold immobility in patients
5 after isolated PDA ligation. The presence of symptoms such as dysphonia (48%), dysphagia
6 (27%), enteral tube feeds (24%) and respiratory symptoms i.e., stridor, increased work of
7 breathing (11%) were observed in this patient cohort with a median follow-up of 3 years.
8 This paper concluded that there is an association between the severity of clinical
9 dysfunction at presentation and clinical outcomes [53].
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13 Karas et al. [11] advocate that parents and primary care providers should be counselled that
14 there is a 46% probability of children requiring a surgical feeding tube after sustaining VCP
15 post-cardiac surgery. They also discuss, in addition to others [54, 55, 56] that infants
16 undergoing correction of larger structural defects with longer operative times and more
17 extensive dissection near the recurrent laryngeal nerve have shown greater rates of
18 intubations which is speculated to be due to lesser contralateral vocal fold compensation
19 [17].
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23 **5. Quality of life (QoL):**

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25 Considering the patient's QoL is crucial to better understand the consequences of VCP and
26 its treatment [57]. Despite a catalogue of available instruments for its assessment, QoL
27 remains a complex concept that is defined and interpreted in a variety of ways across and
28 within various medical disciplines globally [57]. The utility of the Paediatric Voice-Related
29 Quality of Life (pVRQOL) survey was validated in a study by Boseley et al. [58] as a more
30 comprehensive instrument than the Paediatric Voice Outcomes Survey (pVOS). Whilst
31 pVRQOL results report social-emotional and physical-functional effects, pVOS is less
32 sensitive and only documents global preoperative and postoperative changes [58]. Walz et
33 al. [59] found that the pVRQOL score in premature paediatric patients with a history of
34 cardiac surgery is significantly lower than those without a history of cardiac surgery. There
35 was no explanation given as to whether the reason for a decrease in the pVRQOL score was
36 due to recurrent laryngeal nerve injury. To our knowledge, there is no literature that directly
37 associates QoL, recurrent laryngeal nerve injury (and hence VCP), and paediatric cardiac
38 surgery. All references to QoL are thus noted to be anecdotal in nature.
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42 Post-cardiac surgery, it is clear that children are at significant risk of aspiration (which could
43 lead to pulmonary injury) and dysphagia due to vocal fold dysfunction [54, 60, 52]. Reports
44 documenting post-operative occurrences are abundant yet there is a scarcity of data
45 describing long-term follow-up [51, 48, 61]. The effects of dysphonia can be devastating for
46 the social development of a child resulting in shyness, withdrawal at school, and general
47 difficulty in engaging with others [8].
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51 Rodney et al. [8] – whose comparative analysis showed non-significant differences across
52 the majority of postoperative variables between VCP and non-VCP neonates except for
53 aspiration – suggested that the QoL is predominantly impacted when there is insufficient
54 self-resolution of the VCP or when a diagnosis is missed. Follow-up with an otolaryngologist
55 is therefore of major importance yet may be overlooked due to lack of fail-safe institutional
56 measures to ensure that every VCP patient receives a follow-up appointment. This could
57 also be amplified further by insufficient foresight from the cardiac team to refer to
58 otolaryngology when noting VCP during their designated follow-up [8]. Retrospective
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3 findings from Jabbour et al. [22] revealed only a 24% resolution rate (n=278) in their
4 paediatric cardiac surgery population - which is similar to the 35% (n=80) reported by
5 Truong et al. [51]. They also proposed a regimented follow-up plan that could be adopted
6 for all paediatric VCP patients as shown in figure 1 [22].
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9 Importantly, Rodney et al. [8] highlight that through effective counselling, physicians can
10 fully prepare parents of aortic arch reconstruction patients to both manage aspiration with
11 modified feeding and anticipate potential sequelae. This can also subdue concerns of
12 aspiration-related pneumonia, gastrostomy, and tracheostomy (for BVCP) due to their
13 relative rarity [8]. Resiliency of the neonatal lungs was cited as a potential explanation for
14 the evasion of respiratory sequelae [8].
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18 As described by Dewan et al. [18], in addition to modified feeding, presentations of
19 aspiration and dysphagia may require further dietary augmentation through delivery via
20 gastrostomy or nasogastric tube. They found that VCP neonates are more likely to
21 experience an abnormal cough and gag-reflex and require an increased length of
22 hospitalisation due to the tube feeding [18]. Supportive findings by Sachdeva et al. [55]
23 indicated that 87% (n=38) of VCP paediatric in their cohort were intolerant of full oral feeds
24 at discharge and requiring modification to their nutritional intake, potentially with a
25 modified delivery method. Recent findings from Richter et al. [61] suggest that there is an
26 expected time to regular oral feeding of less than 2 years irrespective of the infants' VCP
27 status and genetic comorbidities. Despite the good prognosis, parents of VCP infants should
28 be informed of modified diets at discharge and that there is a greater long-term risk for
29 inpatient hospitalisation due to feeding difficulties and poor weight gain [61].
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34 Sequelae of VCP such as prolonged tube feeding, and dysphonia could significantly impact
35 both the parent and the child as well as their relationship. More quality-of-life evaluations
36 could be conducted in the future pertaining to the children and their families; voice-related
37 QoL, for instance, is a metric that could be better represented. Whilst Walz. et al. [59]
38 discussed voice-related QoL for a premature paediatric cohort, there could be further
39 pVOS/pVRQOL short-term and long-term studies in a broader-aged paediatric cardiac
40 surgery population for patients experiencing VCP. For older paediatric patients,
41 retrospective QoL studies could be conducted to quantify the extent to which the condition
42 affected their lives prior to intervention and be the substrate for additional future
43 interdisciplinary interventions to better support this group.
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48 **6. Summary:**

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50 Advances in paediatric cardiac surgery and medical care have caused a paradigm shift in
51 patient care and survival. Patients with prematurity and complex congenital cardiac diseases
52 are increasingly surviving their initial challenges and leading productive lives. The possibility
53 of an increased prevalence of VCP in children is therefore likely, particularly with emerging
54 undetected VCP cases. Variation in the reported causes of VCP is likely institution-specific
55 and attributable to the paediatric cardiac surgery activity (figure 2) [6] - it is therefore
56 difficult to ascertain fully representative statistics in regions where healthcare infrastructure
57 remains underdeveloped.
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3 Patients at greater risk of VCP are female, premature and low-birth weight undergoing
4 either or both PDA ligation or aortic arch interventions. Avoiding the serious complications
5 of aspiration and dysphagia are at the forefront of management decisions for the child-
6 surgical feeding tubes are therefore common [11]. Clinicians should be prepared to counsel
7 either or both patients and their families on the QoL implications, which remains an area
8 that requires further study. Certain events, such as aspiration-related pneumonia and
9 gastrostomy, could be informed as rarities for aortic-reconstruction patients [8]. On the
10 contrary, clinicians should expect respiratory complications and enteral feeding for surgical
11 PDA ligation patients [14].
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15 Prospective multi-centre cohort studies are warranted in future and should aim to identify
16 further VCP risk factors and better assess presentations such as voice and swallowing.
17 Longer-term stricter institutional follow-ups will lend to enhanced understanding of true
18 recovery rates. Pre- and post- operative FL should be conducted universally to prevent VCP
19 underestimation in asymptomatic patients and promote early detection. Nonetheless,
20 primary care providers should remain cautious as to not to overlook insidious cases of VCP
21 which many present with similar symptoms to asthma in older children [62].
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28 **7. Conclusion:**

29 Cardiac surgery remains the leading cause of iatrogenic VCP in the paediatric population.
30 Early recognition and management of VCP following cardiac surgery is vital to improve
31 recovery and reduce morbidity and mortality. This is particularly important given that VCP
32 can present insidiously or be mistaken for other conditions such as asthma. There are
33 limited studies looking at VCP post cardiac surgery, making it difficult to accurately assess
34 and evaluate its implications, and translate findings into clinical practice. Follow-up studies
35 targeting this cohort are warranted.
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For Peer Review

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Tables:

	Injection Laryngoplasty	Type 1 Thyroplasty	Non-selective recurrent laryngeal reinnervation
Advantages	<ul style="list-style-type: none"> -Immediate improvement -Minimally invasive (endoscopic, outpatient procedure) -Minimal anaesthetic use 	<ul style="list-style-type: none"> -Immediate improvement -Reversible - can remove implants. -Less immunogenicity was reported for materials than for IL. 	<ul style="list-style-type: none"> -Preserved laryngeal muscle bulk -Does not preclude medialisation procedures -Minimal foreign body and inflammatory reaction risk. -Greater posterior glottic closure - no adjunct surgery needed. -General anaesthesia - does not require patient cooperativity [35]
Disadvantages	<ul style="list-style-type: none"> -Limited duration - ideal materials last up to a few months. -Irritation to native tissue and histopathological changes. -Reduces vocal fold elasticity - weaker voice 	<ul style="list-style-type: none"> -Limited duration - requires revisions to accommodate developing anatomy. -Laryngeal muscle atrophy - progressive denervation -Pre-adolescent paediatric patients pose procedural challenges -Procedure is normally done awake or under light sedation and requires cooperativity. -General anaesthesia may be required for which concerns exist about neurocognitive effects with prolonged procedures [34] Involving children under 3 years old [37]. -Requires adjunct arytenoid adduction because it does not correct persistent posterior phonatory gaps 	<ul style="list-style-type: none"> -Average time of 4.5 months to improve voice quality [33] - observation period in older children is ~12 months. -General anaesthesia - neurocognitive impairment [41] risk if under 3 years old [37]

Table 1: Comparison of the positives and drawbacks of injection laryngoplasty, Type 1 thyroplasty, and NSLR adapted from Espinosa et al [21] unless indicated otherwise.

Author	Year of publication	Surgery	Incidence	Recovery	Median time of follow up (months)
Karas et al. [11]	2019	Cardiac surgery	100% (65/65)	29% (19/65)	4.3
Pourmoghadam et al. [49]	2017	Aortic arch repair by either:	48% (43/89)	Overall recovery not specified	11
		Norwood repair	41% (26/43)	74% (14/19)	
		Non-Norwood repair	65% (17/43)	86% (12/14)	
Alfares et al. [10]	2016	Congenital cardiac operations including aortic operation, PDA ligation, Norwood repair, and median sternotomy	1.1 % (32/3036)	61% (19/32)	10 months for those who had recovered
Jabbour et al. [22]	2014	Cardiac surgery	68.8% (278/404)	24% (68/278)	Not stated
Truong et al. [51]	2007	Congenital cardiac operations (PDA ligation and complex cardiac procedure)	100% (109/109)	35% (25/80)	16.4
Pereira et al. [56]	2006	PDA ligation	11.5% (7/61)	71% (5/7)	9
Khariwala et al. [60]	2005	Congenital cardiac operations	Unilateral – 19% (11/48)	82% (9/11)	3

			Bilateral – 6% (3/48)	Not stated	Not stated
Zbar et al. [52]	1996	PDA ligation	8.8%, 6/68	0%	6
Fan et al. [50]	1989	PDA ligation	4.2% (7/167)	0%	Not stated

Table 2: A summary on the incidence, recovery and follow up period for VCP following cardiac surgery [10,11,22,49,50,51,52,56,60]

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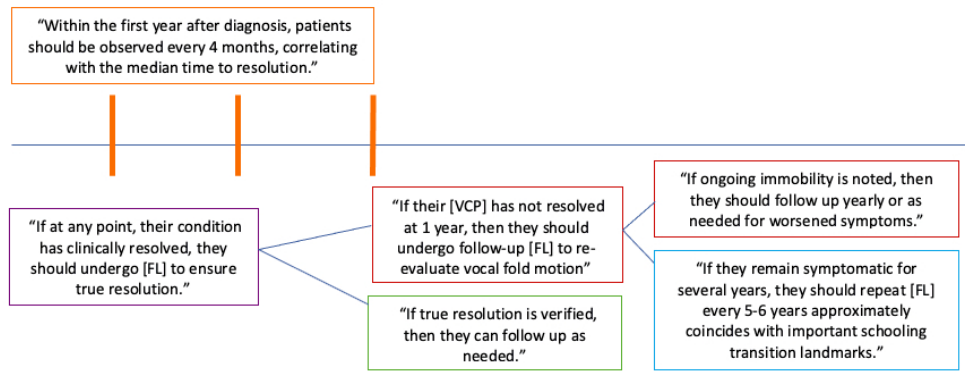


Figure 1: A flow chart summarising the recommended otolaryngology follow-up algorithm adapted from Jabbour et al. [22]

Figure 1. classification of risks of each surgical procedure in relation to developing vocal cord palsy
311x143mm (72 x 72 DPI)

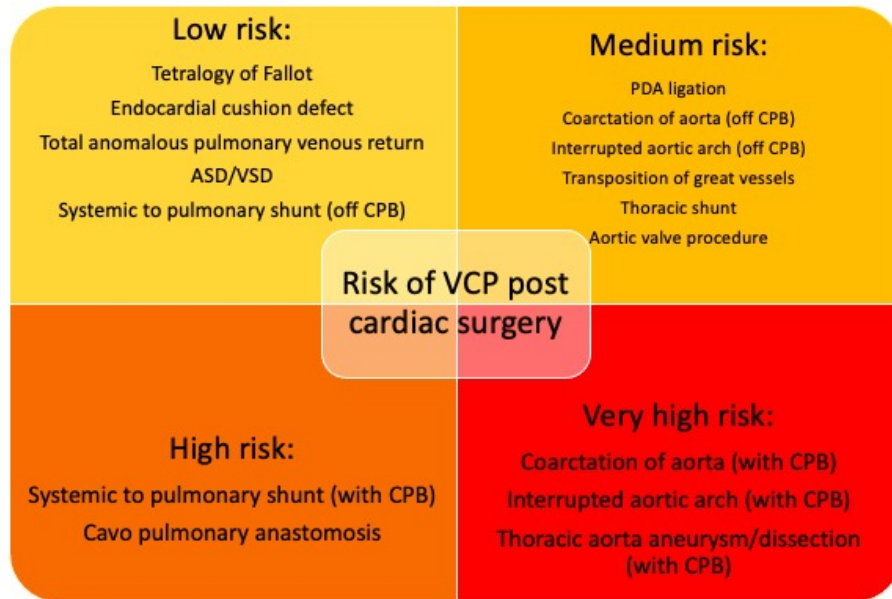


figure 2

224x155mm (72 x 72 DPI)