ORIGINAL ARTICLE

Artificial Lung / ECMO



Rescue extracorporeal membrane oxygenation for massive anterior mediastinal masses

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Received: 10 June 2020 / Accepted: 25 March 2021 / Published online: 3 April 2021 © The Japanese Society for Artificial Organs 2021

Abstract

The management of massive anterior mediastinal masses (AMM) is challenging. With the burgeoning role of extracorporeal membrane oxygenation support (ECMO) beyond the confines of salvage therapy, more trained clinicians are adopting it as a bridge for high-risk procedures or situations where temporary respiratory or cardiac support is required. We report our experience with using ECMO in the management of massive AMM in this case series of three patients sharing their clinical details and the lessons learned from them.

Keywords ECMO · Anterior mediastinal mass · Mediastinal mass

Introduction

The management of massive anterior mediastinal masses (AMM) poses both a diagnostic and therapeutic challenge. The symptoms and signs vary from patients being asymptomatic to presenting in extremis secondary to compression on mediastinal structures. The superior vena cava (SVC), pulmonary arteries (PA), and trachea are compressed in that order because of their inherent wall tension. Due to the slow-growing nature of AMMs, patients experience symptoms insidiously. Radiological diagnosis of AMM is easily made with computed tomography (CT) scans but the histological diagnosis that further guides treatment entails patients undergoing a biopsy. The anaesthetic considerations

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in patients with AMM are complex and have been widely discussed [1]. Extracorporeal membrane oxygenation (ECMO) has been described in reports as a successful means of bridging patients with AMMs causing extrinsic airway compression during induction chemotherapy or surgical treatment [2–15]. We present a series of patients who underwent Veno-arterial (VA) ECMO for management of their AMM and review literature on the use of this modality. We share our threshold for initiating ECMO in patients with massive AMM and encourage early adoption of ECMO to improve survival.

Case series

The study has obtained local DSRB approval (2018/01257). Waiver of consent was given.

Case 1

A 40-year-old lady with a large left goitre with retrosternal extension presented with compressive symptoms of dysphagia, shortness of breath, and stridor. She was unable to lie flat and slept prone. Due to her symptoms, surgical excision was offered. CT thorax showed a 11.6×11.0 cm left thyroid lobe nodule with mass effect displacing the left sternocleidomastoid muscle, internal jugular vein, common



Fig.1 CT Thorax showing large goitre (*) with displacement of carotid vessels (white arrow) and compression of trachea (black arrow)

carotid artery and severe narrowing of the trachea (Fig. 1). In view of the airway risk, ECMO support was activated for the left hemithyroidectomy. A multidisciplinary discussion was held where it was discussed that in the event of failed intubation, veno-venous (VV) ECMO would be initiated. The patient could not be intubated despite attempts at awake fibreoptic intubation. Thus VV ECMO support was started in lieu of ventilation. Bilateral femoral veins were cannulated by open cut down, in view of the patient's large body habitus (body mass index 34.6), under local anaesthesia. 3.5 L/min of flows were achieved with the drainage cannula in the inferior vena cava and the return cannula in the right atrium. The cannulas were positioned appropriately so as to exclude recirculation. However, this was inadequate to maintain oxygenation and the cannulation strategy was 451



Fig. 2 CT Thorax showing AMM (*) compressing trachea (black arrow) and obliterating the SVC and brachiceophalic vein

changed to femoral VV-A ECMO. This permitted higher flows of 5.5 L/min but the patient had differential hypoxemia with right upper limb saturation remaining at 60%. A semi emergent tracheostomy was performed which enabled better ventilation and improved the saturations to above 90%. The left thyroidectomy was then performed under ECMO support and the patient was weaned off ECMO post-operatively. Post-operatively patient recovered and her tracheostomy decannulated prior to discharge on day 24.

Case 2

A 23-year-old gentleman presented with worsening shortness of breath, stridor, and hoarseness of voice. He was diagnosed 6 months prior with germ cell tumour of the mediastinum but defaulted follow-up. CT scan then showed a 14.0×10.0 cm AMM with SVC obstruction, PA and airway compromise. A repeat CT after 6 months showed that the mass had increased to 16.0×13.0 cm with severe compression of great vessels, airway, and left phrenic nerve causing diaphragmatic palsy (Fig. 2). Due to his impending airway compromise, his right femoral artery and vein were percutaneously cannulated with a 7Fr sheath in preparation for ECMO. He underwent awake fibreoptic intubation with a size 8.5 armoured cuff endotracheal tube (ETT) and bronchoscopy in the operating theatre. Distal to the ETT, the carina was compressed 80% in diameter, the left main bronchus compressed 70% and right main bronchus compressed 90%. The distal bronchi were patent. The patient was transferred to the intensive care unit (ICU) after intubation but remained unstable with difficulties in ventilation. The ETT was adjusted under bronchoscopic guidance beyond the collapsed part of the trachea but the patient deteriorated and had a < 1 min episode of pulseless electrical activity arrest. He was resuscitated but required inotropic support (Noradrenaline 0.1 mcg/kg/min) and had poor gas exchange. In view of the haemodynamic instability and possibility of worsening tumour size after initiation of chemotherapy, decision was made to stabilise the patient on VA ECMO. VA ECMO was initiated via the cannulated femoral vessels. Chemotherapy was started and serial CTs showed a decrease in the size of the AMM. He was decannulated on day 8. However, he failed extubation due to tracheal and bronchomalacia for which he underwent both tracheal and bronchial stent insertion. He was eventually extubated on day 39. As a result of immunosuppression, he sustained an infected pseudoaneurysm of his femoral artery which required repair with a pericardial patch and sartorius muscle flap. The patient was discharged from the hospital on day 58.

Case 3

A 29-year-old gentleman presented with progressive facial swelling over a month associated with shortness of breath, productive cough, orthopnoea, and unintentional weight loss. CT Thorax showed a 14.1×11.7 cm AMM causing SVC obstruction, compression of bilateral brachiocephalic trunks, thrombosis of the right internal jugular vein, and distortion of the main and bilateral PAs (Fig. 3). He underwent bilateral anterior mediastinotomy and biopsy of AMM with ECMO standby support. The left femoral artery and vein were percutaneously cannulated with 7Fr sheaths. During the procedure, the patient desaturated was intubated and initiated on femoral VA ECMO support. The surgical biopsy was completed and he was shifted to ICU for further management. Histology revealed primary mediastinal B-cell lymphoma. He was initiated on chemotherapy and intravenous steroids with prophylaxis for tumour lysis syndrome. The tumour size did not change despite two cycles until he received a fraction of radiotherapy. He was decannulated after 12 days of VA ECMO. Serial CTs showed improvement in mass effect on the trachea and PAs. He had difficulty weaning off the ventilator requiring multiple bronchoscopies to evacuate bronchial clots and secretions to open up the distal bronchi. His left mediastinotomy wound became infected with Elizabethkingia requiring VAC dressing and Levofloxacin. The patient eventually recovered and was discharged home on day 140.

Discussion

We share our experience of three patients with AMM who successfully had ECMO for hemodynamic support as part of their diagnostic and therapeutic management. The mortality of two of our patients prompted us to recalibrate our approach towards earlier ECMO team activation and create a workflow for high-risk patients with AMM (Table 1). We observed that delayed diagnosis of massive AMM, late

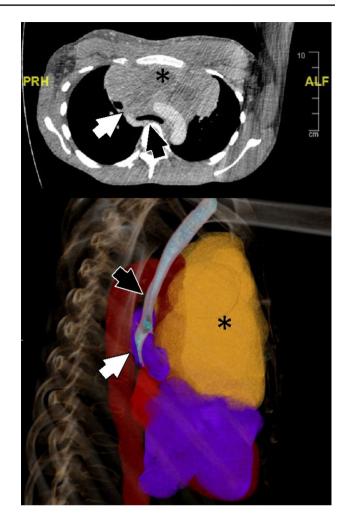


Fig. 3 CT Thorax and 3D reconstruction showing AMM (*) causing tracheal compression (black arrow) and SVC compression (white arrow)

referrals for ECMO, and lack of preparedness contributed to mortality. Our first mortality (MABS) was in a, young patient who collapsed in the haematology procedure room when he was sedated and positioned for a bone marrow aspirate in the right lateral position. When patients with massive AMM are deeply sedated or paralyzed, the weight of the AMM causes acute airway or great vessel occlusion precipitating cardiopulmonary compromise. Cardiopulmonary resuscitation was started and the decision for ECMO activation was delayed. By the time ECMO was initiated, it may have been too late and patient's heart and brain had infarcted. Our second patient (MSBR), who died, had an open biopsy in the operating theatre with anaesthesia support. Unfortunately, massive tumour bleeding was encountered during the procedure which rapidly expanded the tumour and exacerbated the compressive symptoms. While the call for ECMO initiation was made expediently, access lines were not yet placed and the team had to be called in during out-of-office hours. This

Table 1	Summary	of cases	with	massive	AMM	requiring ECMO)
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Patient	MABS	MSBR	CSL	BH	NHL
Age	19	23	40	23	29
Gender	Male	Male	Female	Male	Male
Weight (kg)	60	101	100	100	60
Height (cm)	170	174	170	180	164
BSA	1.70	2.15	2.10	2.20	1.65
Size of mass (cm)	12.0×8.8	13.2×10.6	11.6×11.0	16.0×13.0	14.1×11.7
Compression of airway	Yes	Yes	Yes	Yes	Yes
Compression of SVC	No	Yes	No	Yes	Yes
Compression of great arteries	Yes	No	No	Yes	Yes
Compression of phrenic nerve	No	No	No	Yes	No
Bleeding	Yes	No	No	No	No
Etiology of mass	Acute lymphoblastic leukaemia	Diffuse large B cell lymphoma	Retrosternal goitre	Germ cell tumour	Primary mediastinal B-cell lymphoma
Surgery	Bone marrow aspirate	Chamberlain under LA convert emer- gency right hemi- clamshell	Left hemithyroidec- tomy	Awake fibreoptic intubation	Bilateral chamberlair under LA
ECMO type	VA ECMO	VA ECMO	VV to VVA ECMO	VA ECMO	VA ECMO
Cannulation strategy	Percutaneous RCFV then cutdown RCFA	Right groin cutdown for RCFA + RCFV	Bilateral groin cutdown, bilateral femoral veins then RCFA	Right groin cutdown for RCFA + RCFV	Right groin cutdown for RCFA + RCFV
Duration of ECMO run (days)	1	1	1	8	12
Weaned off ECMO	No	No	Yes	Yes	Yes
Inpatient mortality	Yes	Yes	No	No	No
On ECMO mortality	Yes	Yes	No	No	No
ECMO site infection	No	No	No	Yes	Yes

delay contributed to the patient's eventual demise. Through multiple root cause analyses and multidisciplinary discussions, we formulated a protocol (Fig. 4) which involved early consultation and involvement by oncologists, intensivists, radiologists, anaesthetists, and cardiothoracic surgeons once a patient is diagnosed with an AMM. Patients identified as high-risk would undergo surgical biopsy in the operating theatre with anaesthesia and standby ECMO support. Femoral access sheaths are inserted under local anaesthesia prior to biopsy to expedite ECMO initiation when necessary. A primed ECMO circuit is set up in the theatre as the case is underway. The patient undergoes awake fibreoptic intubation, if needed, to secure the airway. Should intubation fail or patient develop peri-arrest symptoms, ECMO is immediately instituted with cannulation under local anaesthesia. After the biopsy, the patient is monitored in a high dependency unit. If no cannulation occurs, the groin lines are kept postoperatively for 24 h to ensure the patient tolerates induction therapy. Thirteen patients were referred after the protocol was devised of which the three patients in this series went on ECMO.

There is an increasing role of ECMO beyond the confines of salvage therapy as a bridge for temporary cardiopulmonary stabilization. This can be applied to patients deemed high risk for cardiac arrest when undergoing procedures. For patients with AMM, these include anterior mediastinotomy or radiological guided core biopsies. ECMO use has been well described for patients with lymphoma requiring invasive biopsy for histology [2-6]. More recently, ECMO has been used intraoperatively for cardiopulmonary support of patients with anticipated difficulties of securing airway or ventilation [7]. The use of ECMO in a subtotal thyroidectomy with distorted airway as a means for oxygenation has been reported [8]. Others have also reported the use of ECMO for the duration of surgery when resecting mediastinal and bronchoscopic masses [9–11]. Placing the patient on ECMO eliminates the need for constant airway ventilation during surgery allowing for better oncological resection.

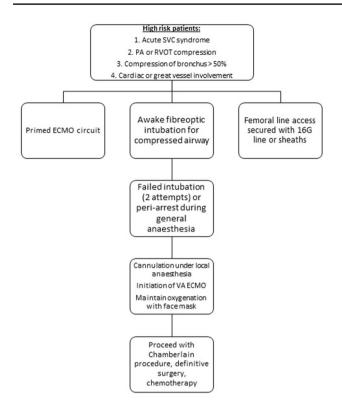


Fig.4 Simplified mediastinal mass protocol and indications for ECMO

Beyond these procedures, ECMO has also been used in a supportive role for patients with AMM compromising cardiopulmonary function. In a reported series of lymphoma patients two of them required ECMO for high-risk mediastinal masses [12]. These patients had severe compression of their PA and SVC which needed stenting on ECMO. Other authors have also reported similar cases of patients needing ECMO for mediastinal masses [13–15]. These cases are summarized in Table 2.

The choice of ECMO mode in such scenarios can be challenging. Several case reports highlight patients with mediastinal masses who were managed either on VV or VA ECMO [8, 9, 14]. VA ECMO would be prudent in patients with extrinsic compression on the airways and vasculature as well as during an emergent peri-arrest setting [8, 14]. Tracheal or bronchial tumours with fixed airway obstruction may be managed on VV ECMO [9]. If in doubt, we recommend insertion of VA ECMO first with a view for an additional venous cannula if required [16]. As seen in case 1, we attempted VV initially but were unable to sustain oxygenation with just venous cannulas. Hence we changed strategy to VVA to improve drainage and thereby provide better flows and improve oxygenation. Patients with AMM have altered neck anatomy or superior venacaval syndrome making additional venous access difficult. In a patient with good cardiac output, this can lead to differential hypoxia, however, it might afford sufficient time for a definitive airway or a tracheostomy to be performed. We encourage open multidisciplinary discussions prior to embarking on such endeavours and to establish clear thresholds not only for instituting but also withdrawing ECMO support [16]. Differential hypoxia should be routinely looked for in all patients with VA ECMO especially if cardiac function remains preserved.

In our experience, the major challenge encountered was that of cannula site infection in these immunocompromised patients. We practice a groin cut down for VA ECMO for better exposure during cannulation, decannulation and repair but this poses a higher infection risk. We mitigate this by cannulating in an aseptic manner, vigilant wound care and treating any potential infection aggressively.

Both percutaneous and cut-down cannulations are viable options depending on a centre's operators' preference and familiarity. Both are shown to have similar outcomes with less infection for percutaneous closure but more vascular complications on decannulation [17]. Percutaneous insertion requires the use of ultrasound image-guided insertion which may be less invasive but requires competency with ultrasound usage. Cut down requires surgical instruments and techniques that non-surgically trained personnel may not be familiar with. Traditionally, both would require femoral artery repair upon decannulation of arterial cannulas. With the advent of perclose devices, percutaneous cannulations may save patients an additional surgery upon decannulation [18]. We recommend that ECMO cannulators adopt the strategy that they are most competent with and prepared for with the necessary equipment available.

Conclusion

Our case series presents adult patients with massive AMM that underwent VA ECMO. We advocate for the use of ECMO in AMM both as a supportive and rescue measure. We believe that as ECMO is further refined and its deleterious effects mitigated, the use of ECMO can be expanded to incorporate wider indications and adopted in a more preventive and elective setting. Our protocol seeks to identify high-risk patients from the outset based on both clinical and radiological criteria. It specifies clearly the multiprong approach that ECMO teams can take in ensuring the safety of patients with AMM.

Acknowledgements We acknowledge the perfusion team at the National University Heart Centre Singapore for their support of our ECMO programme and all other persons involved in the formulation of the workflow.

Author contributions All authors contributed to the study conception and design. Material preparation, data collection, and analysis were

lable 2 Sun	nmary of case	lable 2 Summary of cases in the literature	iture										
Author year	Stewart (1998)	Frey (2006)	Frey (2006) Chao (2006)	Wickiser (2007)	Aboud (2008)	Shao (2009)	Felten (2010)	Oto (2014)	Wohlfarth (2014)	Worku (2015)	Kim (2015)	Lueck (2016)	Nokes (2018)
Number of patients on ECMO	1	-	-	6	_	_	-	-	7	-	-	1	_
Age range	19	10	21	4, 11	43	51	I	40	22-51	41	88	20	49
Gender	Male	Male	Female	Male	Female	Female	Male	Male	8 Male 6 Female	Female	Female	Male	Female
Size of mass (cm)	13×10×5	I	13.6×8.3×16.0	I	1	17×16	23×10×11	12×7	1	13.3	5×3.9	22×12×15	I
Compres- sion of airway	Yes	No	Yes	Yes	Yes	Yes	Yes	Yes	2 Yes	Yes	Yes	Yes	Yes
Compres- sion of SVC	Yes	No	Yes	No	No	No	No	No	I	Yes	No	Yes	No
Compres- sion of great arteries	No	No	Yes	Yes	No	No	No	No	3 Yes	Yes	No	No	No
Bleeding	No	No	No	No	No	No	Yes	No	6 Yes	No	No	No	No
Etiology of mass	Lympho- blastic lym- phoma	Precursor T lympho- blastic lym- phoma	B-cell Burkitt's lymphoma	Lymphoma	Non-Hodg- kin lym- phoma	Multi- nodular thyroid goiter	Mediastinal sarcoma	Precur- sor T lympho- blastic lym- phoma	Lymphoma	Diffuse large B-cell lym- phoma	Malignant teratoma	T lympho- blastic lymphoma	Multiple myeloma and extramed- ullary plasmacy- toma
Surgery	Mediasti- noscopy and biopsy	Biopsy of mediasti- nal mass	Biopsy of medi- astinal mass	Biopsy of mediasti- nal mass, bone marrow aspiration	CT guided biopsy	Subtotal thyroid- ectomy	Resection of Intubation medi- astinal sarcoma	Intubation	1	1	Resec- tion of tracheal mass	1	Endobron- chial stenting
ECMO type	VA	VA	٨٧	VA	٨٧	VA	VA	٧٧	3 VA 11 VV	VA	٧٧	VA	٨٧

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Table 2 (continued)	ntinued)												
Author year	Stewart (1998)	Frey (2006)	Frey (2006) Chao (2006)	Wickiser (2007)	Aboud (2008)	Shao (2009)	Felten (2010)	Oto (2014) Wohlfarth (2014)		Worku (2015)	Kim (2015)	Lueck (2016)	Nokes (2018)
Cannula- tion strategy	Femoral	Cutdown right common carotid and right jugular vein	Bilateral femoral Left f veins ral a righ fem veir	emo- artery, oral	Bilateral femoral veins	. 1	Femoral cutdown	Right jugular vein, Right CFV	VV: right jugular vein, right CFV VA: right CFV, CFA (Right subcla- vian patch)	Bilateral femoral veins	Bilateral femoral veins	Bilateral femoral vessels	Bilateral femoral veins
Duration of 2 ECMO run (days)	7	9	ε	4, 6	9	Ч	1	9	3–34	S	140 min	7	1
Weaned off Yes ECMO	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	7 Yes 7 No Yes	Yes	Yes	Yes	Yes
On ECMO No mortality	No	No	No	No	No	No	No	No	7 Yes	No	No	No	No

performed by LL, HKS, KYJ, LT, HM, and KR. The first draft of the manuscript was written by LL and all authors commented on previous versions of the manuscript. All authors read and approved the final manuscript.

Funding None.

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