

REFERENCES

1. Has C, Sparta G, Kiritsi D, Weibel L, Moeller A, Vega-Warner V, *et al.* Integrin alpha3 mutations with kidney, lung, and skin disease. *N Engl J Med.* 2012;366:1508-14.
2. Nicolaou N, Margadant C, Kevelam SH, Lilien MR, Oosterveld MJ, Kreft M, *et al.* Gain of glycosylation in integrin alpha3 causes lung disease and nephrotic syndrome. *J Clin Invest.* 2012;122:4375-87.
3. Yalcin EG, He Y, Orhan D, Pazzagli C, Emiralioğlu N, Has C. Crucial role of posttranslational 79 modifications of integrin alpha3 in interstitial lung disease and nephrotic syndrome. *Hum Mol Genet.* 2015;24:3679-88.
4. He Y, Balasubramanian M, Humphreys N, Waruiru C, Brauner M, Kohlhase J, *et al.* Intronic ITGA3 mutation impacts splicing regulation and causes interstitial lung disease, nephrotic syndrome, and epidermolysis bullosa. *J Invest Dermatol.* 2016;136:1056-9.
5. Colombo EA, Spaccini L, Volpi L, Negri G, Cittaro D, Lazarevic D, *et al.* Viable phenotype of ILNEB syndrome without nephrotic impairment in siblings heterozygous for unreported integrin alpha3 86 mutations. *Orphanet J Rare Dis.* 2016;11:136.
6. Cohen-Barak E, Danial-Farran N, Khayat M, Chervinsky E, Nevet JM, Ziv M, *et al.* A nonjunctional, nonsyndromic case of junctional epidermolysis bullosa with renal and respiratory involvement. *JAMA Dermatol.* 2019; 155:498-500.

Successful Right Atrium-Pulmonary Artery ECMO in an Infant With Severe Necrotizing Pneumonia and Bilateral Bronchopleural Fistula

We report an infant with necrotizing pneumonia and bilateral broncho pleural fistula, who failed on conventional and high frequency ventilation and was managed successfully on Venovenous Extra Corporeal Membrane Oxygenator (V-V ECMO) with a unique configuration for 12 days, and weaned off successfully.

Keywords: Severe Pneumonia, Management, Ventilation.

Necrotizing pneumonia is a rare form of complicated pneumonia, which often develops secondary to organisms like *Staphylococcus aureus* and *Streptococcus pneumoniae*. It is often difficult to manage, and occasionally develops complications like pneumothorax and broncho-pleural fistula (BPF).

An 11-month-old infant, weighing 9 kg, was referred to us for respiratory distress and fever of 3 days. He had no significant past medical history, was immunized for age and thriving well. Clinical examination revealed features of bronchopneumonia. He was started on high flow nasal cannula (HFNC) oxygen and broad-spectrum antibiotics. Investigations revealed pancytopenia with elevated inflammatory markers. His clinical condition worsened on day 3 of admission with increasing oxygen requirement (FiO₂ 60%) and High flow nasal cannula (HFNC) support (flow 20L/min). He was electively ventilated and put on pressure regulated volume control (PRVC) mode of ventilation. Ventilation protocol was set to targets as suggested in Pediatric acute respiratory syndrome (pARDS) guidelines. The PaO₂/FiO₂ (PF ratio) was less

than 150 with PCO₂ <70mm Hg and PH >7.2. Initial blood culture grew *Psuedomonas* and injectable meropenam treatment was started. Prone ventilation was tried but had to be discontinued after two hours as saturations worsened. X-ray showed multiple large pneumatoceles. On day 7, he developed pneumothorax on right side; draining intercostal tube showed continuous bubbling suggesting a bronchopleural fistula (BPF). Ventilation was continued with high frequency oscillatory ventilation (HFOV). Maximum settings on HFOV were FiO₂ of 60%, mean airway pressure (MAP) of 16 cm H₂O and amplitude (delta P) of 45. Child did show some improvement with improving blood gases and was maintained on neuro-paralysis. After one week on HFOV, X-ray showed regressing pneumatoceles with PF ratio improving to >200. PaCO₂ was consistently less than 60 mm Hg and PH >7.3 with HFOV settings of FiO₂ 40%, MAP of 14 cm H₂O and amplitude of 40. On day 13, he was weaned off paralysis and changed back to PRVC mode, but developed tension pneumothorax on left side on the next day.

Considering the child to have refractory Acute respiratory distress syndrome (ARDS) with bilateral air leak, Venovenous ECMO was initiated on day 15 of hospital admission. Patient's jugular vein on ultrasound was found to be small hence we decided for central open ECMO. Child was initially put on Venovenous configuration with inflow and outflow cannulas in right atrium, but had to be re-configured in view of poor flow and re-circulation. Right atrium was cannulated with 22F cannula and pulmonary artery with 14 F cannula, and flow of 900-1000 mL/min was obtained. He was maintained on ECMO for 12 days. On day 12 of ECMO, prior to weaning a bronchoscopic clearance and lavage was taken, which showed carbapenam resistant *Acinetobacter* on culture. Antibiotics were accordingly changed to colistin and tigecycline. Post-ECMO weaning on ventilator, child did

not have air leak on either side and intercostal tubes were removed. He was successfully discharged after 55 days of hospital stay. Chest X-ray before discharge showed near total resolution of pneumatoceles.

The exact pathogenesis of necrotizing pneumonia is not completely understood [1,2]. Necrotic areas may give way resulting in pneumothorax or BPF. When air leak develops the ventilation often becomes challenging. Ensuring acceptable gas exchange with minimum added barotrauma from ventilation is essential. Bilateral broncho-pleural fistula has a mortality risk of 20-50% [3]. Split lung ventilation or differential lung ventilation has been described in necrotizing pneumonia with unilateral broncho-pleural fistula. Several other strategies include endobronchial plugging with human fibrin glue in small fistulas [4], autologous pleural patch, video-assisted thoracoscopic fistulectomy or stapling and pneumonectomy are described in selected cases [5]. Our patient had limited options, as the disease was bilateral and extensive with multiple necrotic areas. High frequency oscillatory ventilation (HFOV) is an option for refractory ARDS and air leaks who fail on conventional ventilation [6]. HFOV eliminates the 'inflation-deflation' cycle. It maintains gas exchange with very low tidal volume and optimum mean airway pressure, but this child developed pneumothorax on weaning back to conventional mode.

Veno-arterial (VA) configuration was avoided for ECMO because of higher rates of complications in infants and is not the preferred mode for respiratory failure [7,8]. ECMO cannulation mostly performed in infants is a double lumen internal jugular venous (IJV) cannulation but in our patient IJV lumen was small on screening ultrasound. We initially resorted to an open chest central cannulation with two different cannulas in right atrium. This had to be revised due to poor blood flow and significant re-circulation. We finally tried a unique but less performed, Right atrium-pulmonary artery (RA-PA) configuration to completely eliminate re-circulation.

We have not come across any reports on RA-PA configuration in small children. The main disadvantages of a central open chest ECMO are higher chances of infection and bleeding. The insertion time and bleeding from cannulation site were significantly higher than that described in literature for dual lumen IJV cannula [9]. The case highlights the importance of considering ECMO as a salvage but feasible option in selected cases of severe pneumonia with refractory respiratory failure even in developing countries.

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REFERENCES

1. Hoppe P-A, Holzhauser S, Lala B, Bühner C, Gratopp A, Hanitsch LG, *et al.* Severe infections of panton-valentine leukocidin positive Staphylococcus aureus in children. *Medicine (Baltimore)*. 2019;98:e17185.
2. Masters IB, Isles AF, Grimwood K. Necrotizing pneumonia: An emerging problem in children? *Pneumonia (Nathan)*. 2017;9:11.
3. Alohalı AF, Abu-Daff S, Alao K, Almaani M. Ventilator management of bronchopleural fistula secondary to methicillin-resistant staphylococcus aureus necrotizing pneumonia in a pregnant patient with systemic lupus erythematosus. *Case Rep Med*. 2017;2017:1492910.
4. Goussard P, Gie RP, Kling S, Kritzingner FE, Wyk J van, Janson J, *et al.* Fibrin glue closure of persistent bronchopleural fistula following pneumonectomy for post-tuberculosis bronchiectasis. *Pediatric Pulmonol*. 2008;43:721-5.
5. Gerdung CA, Ross BC, Dicken BJ, Bjornson CL. Pneumonectomy in a child with multilobar pneumatocele secondary to necrotizing pneumonia: Case report and review of the literature. *Case Rep Pediatr*. 2019;2019:2464390.
6. Meyers M, Rodrigues N, Ari A. High-frequency oscillatory ventilation: A narrative review. *Can J Respir Ther*. 2019;55:40-6.
7. Kovler ML, Garcia AV, Beckman RM, Salazar JH, Vacek J, Many BT, *et al.* Conversion from venovenous to venoarterial extracorporeal membrane oxygenation is associated with increased mortality in children. *J Surg Res*. 2019;244:389-94.
8. Ham PB, Hwang B, Wise LJ, Walters KC, Pipkin WL, Howell CG, *et al.* Venovenous extracorporeal membrane oxygenation in pediatric respiratory failure. *Am Surg*. 2016;82:787-8.
9. Moscatelli A, Buratti S, Gregoretta C, Lampugnani E, Salvati P, Marasini M, *et al.* Emergency percutaneous, bicaval double-lumen, ECMO cannulation in neonates and infants: Insights from three consecutive cases. *Int J Artif Organs*. 2015;38:517-21.