Interdisciplinary Peripartum Management of Acute Respiratory Distress Syndrome with Extracorporeal Membrane Oxygenation – a Case Report and Literature Review

Extrakorporale Membranoxygenierung für das interdisziplinäre peripartale Management von schwerem akutem Atemnotsyndrom – ein Fallbericht mit Literaturübersicht

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Abstract
Extracorporeal membrane oxygenation (ECMO) is increasingly used for the management of acute severe cardiac and respiratory failure. One of the indications is acute respiratory distress syndrome (ARDS) for which, in some severe cases, ECMO represents the only possibility to save lives. We report on the successful long-term use of ECMO in a postpartum patient with recurrent pulmonary decompensation after peripartum uterine rupture with extensive blood loss.

Zusammenfassung

Introduction
Obstetrical patients rarely end up at intensive care units (ICU). Excessive postpartal bleeding and hypertensive disorders are the most common reasons for ICU admissions. Almost 10 to 15% of ICU patients and up to 20% of mechanically ventilated patients meet criteria for acute respiratory distress syndrome (ARDS) [1]. Large blood loss and pre-existing bad pulmonary conditions represent two risk factors for ARDS. This acute, diffuse, inflammatory lung injury often ends up in long-term intensive care stays. Numerous other complications may follow. We describe the outcome and emerging problems of a complicated delivery with postpartal ARDS which needed emergency admission to ICU and a long-term therapy with extracorporeal membrane oxygenation (ECMO) for a total of 50 days.

Case Report
A 34-year old female patient from Somalia (gravidity 6, para 5) presented at 38 weeks of gestation at our hospital for birth registration. She reported three vaginal deliveries in Somalia and had two Caesarean sections (CS) in Germany. She roughly mentioned "lung problems" in her medical past which were supposed to be resolved. Her exercise capacity seemed unaffected. Exact details of the medical history and previous operations were blurred by the patient (no clear information about which hospitals had been involved were given). The ultrasound examination showed a normal fetus growth with normal Doppler-findings and a normal amniotic fluid index. During the general examination she presented signs of preeclampsia (increased blood pressure (> 160/100 mmHg), protein/creatinine ratio of 900 mg/g). Therefore the medical indication for induction of labour was explained to the patient. Partly due to religious convictions the patient planned to have more children in the future, so spontaneous labour instead of third Caesarean section was discussed. Aware of increased risk for uterine rupture the patient decided for spontaneous delivery and gave her informed consent. The vaginal examination showed no female genital mutilation, a soft cervix, approximately 1 cm long and with a uterine orifice open to 2 cm, the child’s head fixed. First an induction trial with an
extraamniotic Foley balloon catheter was made. As the balloon did not stay in place however, 1 mg of prostaglandin gel was used intravaginally at Bishopscore 5. During the first stage of labour at 3 cm cervical dilatation a pro-
lapse of the child’s arm occurred. While the patient was getting 
prepared for CS fetal bradycardia lead into emergency CS. Intra-
operatively a large rupture of the anterior uterine wall became 
apparent. After safe delivery of an initially limp baby (Apgar 2/ 
10/10, pH 6.99) with direct transfer to the paediatricians this 
rupture could be sutured with single stitches in the area of utero-
tomia and finally with 2 layers of continuous stitches. Due to ex-
cessive intrapartum bleeding (haemoglobin 5.4 mg/dl, blood loss 
approximately 1.5 litres) the patient was transferred to intensive 
care unit (ICU). There she was stabilized with blood transfusion. 
A perioperative antibiotic prophylaxis (cefuroxime) was initi-
ated. The patient was transferred to the obstetric ward 24 hours 
after CS. On day 2 after CS the patient suffered more and more 
from pronounced dyspnea. The cardiologic examination showed 
normal results and pulmonary embolism was excluded by CT 
scan. However, the CT showed bipulmonary infiltrates and shadow-
ons. On suspicion of pneumonia the patient was treated with a 
broad-spectrum antibiotic (imipenem). In the further course her 
general condition worsened so that at one point when the pa-
tient suffered from massive dyspnea the resuscitation team was 
alerted. When the team arrived the patient was unconscious 
and with low heart rate which became an asystolia during the 
intubation procedure and lead to resuscitation. CPR was per-
formed for 2 minutes during the intubation procedure. Satura-
tion at this time was not documented. The blood gas analysis 
afterwards showed pH 7.375, pO2 171.9 mmHg, BE 2.2 and lac-
tate 0.9 mmol/l. Again we admitted the patient to ICU at day 5 
after CS.
During the following days the patient underwent lung protective 
ventilation with low tidal volumes (4–6 ml/kgBW) in relation to 
the patient’s ideal body weight, plateau pressures were below 
30 cm² H₂O. PEEP was adjusted between 10 and 16 cm² H₂O. Prone 
positioning of the patient was tried several times, but oxy-
genation was not improved (non responder/recruiter). Prone 
positioning was kept for 14–16 h; during that period the patient 
required four cardiopulmonary resuscitations due to recurrent 
bradycardia of unknown origin. Bradycardia and asystolia occurred 
reflextory since the first day in our ICU for example after cough-
ing of the patient, after mouth care or when changing tubus po-
tation. The patient was discharged home after 115 days of ICU 
treatment without cardiopulmonary or neurological deficits. 
During her ICU stay, the medical documentation could be pro-
vided. Years ago the patient had undergone multiple pulmonary 
problems such as atypic pneumonia, chronic bronchitis, and a lo-
bectomy of the lower lobe of one lung one year before the deliv-
ery. Furthermore, we found out that our patient required inten-
sive care during her second last pregnancy. A community ac-
quired pneumonia during the third trimester of pregnancy ended 
in ARDS together with renal failure and necessitated intubation, 
dialysis and emergency CS at 34 weeks of gestation. The last preg-
nancy showed a very similar course. After an induction of labour 
because of oligohydramnios the patient developed a restrictive 
ventilation disorder. It ended in secondary Caesarean section at 
40th week of pregnancy and a transfer to ICU for invasive ventila-
tion therapy in total for 11 days again for ARDS of unknown ori-
gin.
The patient was seeking intentionally to mislead us by providing 
false information regarding the aforementioned complications as 
a new pregnancy was not advised after the second last birth due 
to maternal health reasons.
Further family planning was discussed with the patient and her 
husband. We strongly recommended sterilisation or safe contra-
ception. Medical visit at 6 months after discharge was without 
thopathological findings. Mother and baby are in good health.

Discussion

This case reports on four rare complications during labour and 
postpartum period: the occurrence of arm prolapse, uterus rupture 
after two CS, postpartum haemorrhage (PPH) followed by 
ARDS. And finally we report the successful use of ECMO for circu-
latory support and for the treatment of acute respiratory distress 
in a postpartum situation.
Previous studies have reported a general rate of 1/5700 to 1/ 
20000 for uterus rupture in unscarred pregnant uterus [2–4]. 
The overall incidence of uterine rupture in women with a prior 
Caesarean delivery varies between 0.3 to 1 percent [5–7]. A re-
cent large meta-analysis found an incidence of 0.7% for uterus 
rupture/dehiscence in women attempting a vaginal birth after 
one CS (VBAC-1) [7]. Furthermore, several series showed a gener-
al rate of 1.36% of uterine rupture during/after vaginal birth after
two CS (VBAC-2) [8–12]. Although the uterine rupture rate of VBAC-2 is statistically higher than VBAC-1, it should not be forgotten that the absolute rate is very low. This should be kept in mind for patient counselling. In addition, the overall success rate of vaginal delivery after one or two CS is high so that an achievement is possible in 76.5% and 71.1%, respectively [13].

The greatest concern about a trial of labour after a previous CS is a rupture of the scarred uterus. Whereby it should be mentioned that women with induction of labour were more likely to experience uterine rupture or dehiscence than women with spontaneous labour. Therefore, a labour induction indication after CS must be strictly defined. Some factors are important for a high success rate: A previous history of vaginal delivery and a favourable Bishop score are significant. The success rate is between 91% for women with favourable Bishop score and a prior vaginal delivery, and 45% for women with no prior vaginal delivery and an unfavourable Bishop score. For women with no previous vaginal delivery but favourable Bishop score a general rate of 69% successful induction was found [14]. The rupture rate of previously scarred uterus in a large prospective study is 0.4% with spontaneous labour versus 1% with labour induction [15]. These results show that labour induction after CS is possible and clinically justifiable – but, it should only be attempted in appropriately equipped institutions.

ICU admissions during pregnancy and peripartum period have a low incidence of < 1% [16, 17]. Among those patients, the most common causes are PPH and hypertensive disorders. Respiratory failure represents a very rare complication. Differential diagnosis includes pulmonary edema (mostly caused by tocolytic therapy), pneumonia, aspiration, and pulmonary embolism.

In this case, none of the above mentioned pathologies were responsible for the postpartum ARDS. To our opinion, ARDS in this patient was the result of serious tissue injury caused by the uterus rupture with massive blood loss and blood cell transfusion. In our case review the PPH was amplified by the large uterine rupture. The general rate of 1–2.9% of PPH is set in literature [18]. Like in this case serious tissue injury can cause systemic inflammatory response syndrome (SIRS). This is a life threatening post-traumatic complication which is characterized by fever, tachycardia, and tachypnea. These symptoms may resemble a sepsis.

Cellular lesion releases mitochondrial endogenous “damage/danger associated molecular patterns” (DAMPs) that activate the immune system. Mitochondria are evolutionary endosymbions that were derived from bacteria and so bear bacterial molecular motifs. Zhang et al. [19] found that mitochondrial DAMP is released by severe trauma in a similar way that bacterial DNA (PAMPs, pathogen-associated molecular patterns) is released following sepsis. Both use the toll-like receptor-9 (TLR9) to activate neutrophils through activation of p38 MAP kinase (MAPK) enzyme. Similarly, formylated peptides released from bacteria and mitochondria in these settings attract neutrophils by the process of chemotaxis to sites of inflammation and injury through formyl peptide receptor-1 (FPR1) [20]. In both cases, the outcome may be acute lung injury, as part of the SIRS.

On the other hand, the most frequent and severe complication of multiple blood transfusions is the transfusion-related acute lung injury (TRALI). The current pathogenic concepts hold that proinflammatory mediators present in transfused blood products are responsible for the initiation of TRALI [21]. Lee and colleagues [22] hypothesized that mitochondrial (mt)DNA DAMPs are present in blood transfusion products, which may be important in the initiation of TRALI. They extracted DNA from samples of packed red blood cells, fresh frozen plasma (FFP), and platelets procured from their local blood bank. The results showed a range of mtDNA DAMPs in all blood components measured, with FFP displaying the largest variation and they concluded that mtDNA DAMPs are present in packed red blood cells, FFP, and platelets may be mediators of TRALI.

So in this case, the combination of tissue injury by the uterine rupture and the postpartum haemorrhage, which occurs generally in 1 to 5% of deliveries [23], has led to a severe ARDS in a patient with pre-existing bad pulmonary status.

ECMO often represents the last resort for serious acute respiratory distress, although a long-term therapy can lead to life-threatening complications such as haemorrhage and infections. In literature we found little mention of ECMO being used to manage peripartum patients with massive bleeding [24, 25]. Both cases describe the use of ECMO for a short duration (150 respectively 72 hours). In our case the use of ECMO lasted for 50 days which represents an exceptionally long duration compared to literature [24–27].

Others reasons which can lead to ECMO in obstetrical patients are peripartum cardiomyopathy [28], amniotic fluid embolism [29] and transfusion-related acute lung injury [30]. In these cases ECMO lasted 40 to 150 hours and ended each time in a complete recovery of the patient. Main complications of ECMO are bleeding, thromboembolism, cannulation related problems and heparin-induced thrombocytopoenia. Bleeding is the most common complication (30 to 40 percent) of ECMO. In our case even a urological intervention was necessary.

In conclusion, this case report includes an extraordinary sequence of unfortunate circumstances: Beginning with the need of induction of labour due to preeclampsia (with 2 previous Caesarean sections), arm prolapse during delivery in a multiparous patient, fetal bradycardia most likely due to uterine rupture followed by emergency Caesarean section with postpartum haemorrhage. This all followed by deterioration of maternal pulmonary and general condition resulting in ICU admission, consequently several manoeuvres of cardiopulmonary resuscitations and ECMO therapy for almost 2 months. Each of these risk factors is associated with an increased risk of mortality. Nevertheless, our patient could be dismissed in good general condition.

This particular case shows that induction of labour after two prior Caesarean deliveries might be dangerous although newer data showed that labour induction outcomes are similar regardless of whether women have had one or two CS [31]. An early clinical registration of the future mother should be gold standard. It allows obstetricians to identify high-risk pregnancies, take the necessary precautions and might improve the general outcome of mother and child. And finally we can demonstrate that ECMO support for severe respiratory failure in postpartum patients is possible and allows a full recovery even after a long-term therapy with complications.

Conflict of Interest

None.
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