THE COURSE AND OUTCOME OF A PREGNANCY WITH A PREMATURA PRETERM RUPTURE OF MEMBRANES DILEMA OR LIABILITY

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Preterm premature rupture of membranes (PPROM) is in obstetrics with incidence of 0.4% of , with risk for chorioamnionitis 35%, risk for premature deli very very 19%, and sepsis risk of 1%. Primary risk for fetus is pulmonal hypoplasia due to oligohydramnios in 29%, RDS in 66 % of such pregnancies, sepsis in 19%, and contrac ture of extremities in 3%. Fetal death is more than 30% of cases. Patient came to regular pregnancy check up in 22th week of gestation. US exami nation revealed decreased fluid volume AFI 50, with regular morphology and fetal dynamic. She was admitted to hospital. She was given cortico-steroids and antibiotics. In further course of pregnancy amniocentesis was performed in 32th week of preg nancy to rule out the infection and to check pulmonal maturity. In absence of infec tion and lung maturity pregnancy was continued until 36th week of gestation when Cesarean Section was. Baby went well and is now healthy child in the age of three. Some studies suggested that delivery can be significantly prolonged. In our case we maneged to gain 14 weeks, from 22th to 36th gestational week. Firstly gynaelogyst nightmare to ask the woman is she accept therapy. The aim is to deliver a healthy child without neurological and other consequen ces and mother with preserved reproductive function. Controversy still exists when is the appropriate time to finish preg nancy with prolonged PPROM.

Keywords: oligohydramnios, respiratory distress, therapy, outcome.

INTRODUCTION
Preterm premature rupture of membranes (PPROM) is in obstetrics with incidence of 0.4% of , with risk for chorioamnionitis 35%, risk for premature deli very very 19%, and sepsis risk of 1%. Primary risk for fetus is pulmonal hypoplasia due to oligohydramnios in 29%, Respiratory Distress Syndrom(RDS) in 66 % of such pregnancies, sepsis in 19%, and contracture of extremities in 3%. Fetal death is more than 30% of cases.

CASE REPORT
On the 3th of February 2010, a primigravida D. C. (1973) from Belgrade who was in 22th week gestation came to Institut of Gynecology and Obstetrics(IGO) of Clinical Centere of Serbia, Belgrade on a regular check-up. At an ultrasonographic examination a lower level of amniotic fluid AFI – 50 was noticed, with a normal mor-phology and dynamics of the fetus.

On the same day, she was admited to hospital with intensive supervision and with complete laboratory tests (C - reactive protein (CRP), complete blood analysis, wider biochemical parameters: urea, creatinin, lactal dehydrogenesis, alkaline phosphatasis, direct and indi-rect bilirubins, transaminases, cervical and vaginal smears, urin and urino culture.

Before her pregnancy, the patient was diagnosed with hypofunction of thyroid gland so she was already receive ing Tivoral 50 mg/24h and during the pregnancy she was diagnosed with gestational diabetes A1. Microbiological result of cervical and vaginal smear implied bacterial vaginosis and the results of urin showed Escherhia Coli. Antibiotic therapy included amp. Longacef 2g/24h and vaginal tablets Orvagil 1 in the evening with iv. rehydration. The parameters of the infection were monitored daily (CRP and leukocytes with measuring of basal temperature). CRP<5, Le 7.000, but did not go over 36.70C.

Biophysical profile (BPF) was done every day with measuring of amniotic fluid index (AFI) was between 30 and 40 and in the 33 weeks’ gestation it was 15. In the 24th week of gestation with the decision of doctors from Perinatal medicine, the maturation of fetal lungs began with 4mg/6h corticosteroides in the next 24 hours. The parameters of infection were stable.

In the 27th week of gestation a fetal echosonogra phy was performed as a part of a regular perinatal screening and it implied normal echocardiographic finding. Ultrasono graphic result in that period showed normal dynamics of the fetus.
Microbiological result of repeated cervical and vaginal smear showed Klebsielle and Enterobacter in the smears and Escherihia Colli in urine test. According to antibiogram the therapy was Panclav tbl. 625mg/8h and vag. Chloramphenicol a 250 mg in the evening. Control smears sterile, parameters which could show infection - stable. In the 32 week gestation the amniocentesis was performed. The sample was sent to a microbiologi-cal examination and to determine the maturity of fetal lungs. The lectin - phyngomyelin ratio was 1:1,5 and microbiological result showed sterile fetal amniotic fluid.

Obstetric result from the moment of reception and in further course was stable. In the 35 weeks' gestation the patient was sent to doctors at Perinatal medicine who decided to do Cesarean section in the 37 weeks' gestation with intensive monitoring of mother and fetus and intensive monitoring of infection. In the 36 weeks' gestation the patient suffered from contractions and scarce bleeding. Basal temperature was 38,50 C, non-stress test of fetus (NST) implied increase of basal frequency (around 170 a minute). During obstetric examination it was diagnosed that the labour started with first signs of chorioamnionitis. On the same day the Cesarean section was performed and alive, term newborn was born Apgar score (A5) 7/8 with measurements 3600/53/35, on section was performed and alive, term newborn was born.

The baby was admitted to Neonatal Intensive Care Unit and on the fourth day was admitted into semi-intensive care unit. Escherihia Colli was isolated From nasopharynx and eye, which was treated with antibiotics and oxygenotherapy, IV infusion, with a good reaction. The parameters of infection (CRP, Le), after the therapy was included, were in decline. Neurological results: present hypotonia of left side of the body without pathological findings. Important facts: complete blood test and gas analysis within the limits of findings and nuclear magnetic resonance (NMR) of head, torticollis left, luxation of left hip.

The mother was transferred from semi-intensive unit in a good state, afebrile. After 24 hours, increase of basal temperature to 380C was diagnosed. Biochemical microbiological parameters showed presence of infectious agent (Escherihia Colli in uterus smear, lochia and wound), hemoculture - Le - in reference values. Procalcitonin was <0.05 which implied that the patient was not in septic condition.

Antibiotic treatment was included according to the antibiogram: Conet 1g/6h, Vancogal 1g/12h iv infusion, Fraxiparin 0.6 ml/12h. Despite the therapy, the patient had fever in the next 8 days. It led to wound dehiscence which was treated with Amikacin 2g/24h with a good reaction of the patient. In the next few days the parameters of infection (CRP, Se, VFK, Le) were falling and so did the fever. A control ultrasound exa-mination showed that uterus was in good condition, an d consequent follow ups in 3-4 weeks time showed no thing of any ginecological complication.

The baby's condition was intensively monitored at University Children Clinic and at the Institute of Neurology faculty of Medicine at Clinical centre of Serbia, Belgrade. In the 7th month, the baby was diagnosed with luxation of left hip and worked Repositio chirurgica coxae sin. Sec. Ludolf was performed. Neurological result with EEG of head was normal. The child is regulary controlled at University Children Clinic in Tirshova and at Institute of Neurology at Clinical centre of Serbia, Belgrade. The baby is in good overall condition and without consequences. After dismissing from the hospital mother was in the good condition, and consequent follow ups in 2 years time showed no thing of any ginecological complication.

**DISCUSSION**

Premature preterm rupture of membranes (PPROM) prior to fetal viability is a unique and relatively rare problem that is often difficult to manage. It occurs in less than 0.4% of all pregnancies. The major risk is infection, namely chorioamnionitis, which occurs in about 35% of cases. In 19% it causes abruptio placentae in 1% it causes sepsis. [1,2,3,4,5]. The major morbidity in the fetus is lethal pulmonary hypoplasia with oligohydramnios, which occurs in 29% of cases. Other morbidities are respiratory distress syndrome (RDS) (66%), sepsis (19%), IVH (5%), and contractures (3%) [4,5,6,7,8,9]. Fetal death is common and occurs in more than 30%. Older studies have reported that approximately 50% of pregnancies with PPROM deliver in next seven days. More recent studies have shown better prognosis. With appropriate therapy and conservative management, 30% of pre- mature nancies can be prolonged up to 5 weeks [4,5,7,8,10,11,12]. PPROM between 13-26 week of gestation has a poor prognosis although more recent studies have reported encouraging facts. (Waters TP, Mercer BM. The management of preterm premature rupture of the membranes near the limit of fetal viability) [11,12,15,17]. Survival varies with gestational age at diagnosis, from 12% when diagnosed at 16-19 th week, to as much as 60% when diagnosed at 25-26th week. Expectant management may be appropriate in selected patients who are well informed and educated about the risks and the dismal prognosis for the neo-nate [2,4,5,6,7,9,11].

Maternal safety should be the primary concern. Amniocentesis can pro vide information about lung maturity and to give answer of presence of infectious agent in amniotic fluid. This invasive procedure, since the amount of fluid is scant, should be performed by experienced individuals and with obligatory explanation to mother about the potential risks which happen with this procedure [2,4,5,6,7,9,10,11].

The use of corticosteroids to accelerate lung maturi ty should be considered in all patients with PPROM with a risk of infant prematurity from 24-34 weeks' gestation. The use of tocolysis for 48 hours to administer steroids and allow acceleration of fetal lung maturity has been proposed and is being used by some obstetricians. No data support the efficacy of this practice and, as such, when used in this manner, the lack of evidence to support this practice should be discussed with patients to allow informed consent prior to the use of tocolytics and the potential complications and side effects. A positive outcome of pregnancy with premature preterm rupture of membranes is possible with modern peri-natalogical protocols, but still has high risks for mother and for baby [1,2,3,6,7,10,11,12]. The aim of perinatology in these cases is to deliver a healthy baby capable of living with-
out neurological and other sequels, and on the other hand a healthy mother with saved reproductive system. Infection is still a leading risk factor for an unfavourable outcome of these pregnancies because there are more and more kinds of microorganisms resistant to antibiotic therapy (besides modern pharmacotherapy) which largely increases perinatal morbidity and mortality of baby and mother. The initial step in management of PPROM is patient’s consent.

Two of the largest studies that have looked at the efficacy of antibiotic use in PPROM are the National Institute of Child Health and Human Development (NICHD-MFMU) and the ORACLE trial. In the NICHD study, intravenous antibiotics were used for 48 hours - ampicillin 2 g / 6h and erythromycin 250 mg / 6h. The patients were then placed on oral amoxicillin 250 mg / 8h, erythromycin 250 mg / 8h to complete a 7-day course of antibiotic therapy. In this study, the control group, compared with the antibiotic group, had a significantly shorter duration of latency and much greater risk of chorioamnionitis and neonatal sepsis. This therapy refers to beta hemolytic streptococcus group B. The intensive monitoring of mother and fetus is necessary, with daily controls of parameters of infection, weekly cervical and vaginal smear, daily measuring of level of amniotic fluid [3,9,10,11,12].

The maturation of fetal lungs with corticosteroids in these cases is necessary avoid RSD but it is not recommended before 24 weeks’ gestation. The use of corticosteroids in patients with PPROM is solely to accelerate lung maturity but on the other hand it can increase risk of infection. However, current studies in NICHD do not support this concern which is based only on individual studies and meta-analysis. On the other hand, the rates of respiratory distress syndrome (RDS), necrotizing enterocolitis, and fetal intraventricular hemorrhage are decreased. Team work is necessary, including Neonatal unit, and tertiary institutions have a priority in these cases [4,5,6,7,9,10,11].

PPROM in 22 weeks’ gestation with a favourable outcome after 14 weeks is for now a unique case in IGO of Clinical Centre of Serbia, Belgrade and represents the success of the entire team from Institute of Gynecology and Obstetrics of Clinical Centre of Serbia, Belgrade, that is High-Risk Pregnancy Unit, which contributed with their unselfish effort and knowledge to deliver a healthy baby capable for life after 14 weeks with PPROM.

CONCLUSION

Prior to every effort to prolong pregnancy in cases of severe PPROM parents should be in details thoroughly informed about the risks and prognosis for mother and child and give their written consent. We managed to prolong pregnancy after PPROM from 22th to 36th week of pregnancy using antibiotics and corticosteroids and performed Cesarean Section giving birth to healthy child with minor consequences for mother. Child is now 3 years of age and is getting on fine.

REFERENCES

TOK I ISHOD TRUDNOĆE SA PRETERMINSKOM PREVREMENOM RUPTUROM PLODOVIH OVOJAKA DILEMA ILI OBAVEZA

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SAŽETAK
Preterminsko prevremeno pucanje plodovih ovojaka javlja se u manje od 0,4% svih trudnoća, rizik od horioamnionitisa iznosi 35%, od prevremenog porođaja 19%, rizik od seps je 1% . Kod fetusa glavni uzrok morbiditeta je hipoplazija pluća uz pridruženi oligoamnion koji se javlja u 29% slučajeva, respiratorijski distres koji se javlja u 66%, sepsa (19%) i kontraktura (3%) slučajeva. Fetalna smrt je česta i javlja se kod više od 30% slučajeva. Prvorotka, 1973 iz Beograda javlja se 03.02.2010 god. u 22. nedelji gestacije u IGA KCS na redovnu zakazanu kontrolu. Pri UZ pregledu primećena je manja količina plodove vode AFI-50, uz normalnu morfologiju i dinamiku ploda. Trudnica je odmah hospitalizovana i laboratorijski obrađena. Trudnica je dalje vodena antibiotskom terapijom i kortikosteroidima. Porođena je u 36 nedelji gestacije carskim rezom. Ranije studije su ukazivala da se PPRPOM u oko 50% trudnoća završava u narednih sedam dana. Novije studije ukazuju da se porođaj može doživeti odgođiti, ali samo u kliničkim uslovima, uz odgovarajući permanentni stručni nadzor i načelno adekvatne terapije. U našem slučaju trudnica je produžena 14 nedelja (od 22. - 36. nedelje gestacije) uz relativno manje komplikacije. Zadatak akušera u ovakvim slučajevima je dobra procena stanja trudnice i ploda i njena saglasnost da se pristupi terapiji, ali samo u odgovarajućoj ustanovi gde se može pratiti stanje majke i ploda, kako laboratorijski tako i klinički. Cilj je dobiti zdravo i za život sposobno dete bez neuroloških i ostalih sekvela i reproduktivno zdravu majku.

Ključne reči: oligoamnion, horioamnionitis, respiratorijski distres, terapija, ishod.