

Chorioamnionitis - A Serious Perinatal Challenge

Emilija Jasovic-Siveska*

PHO Medihelp, Bitola, North Macedonia, Professor of Gynecology and Obstetrics University St. Clement of Ohrid, Bitola, North Macedonia

***Corresponding Author:** Emilija Jasovic-Siveska, PHO Medihelp, Bitola, North Macedonia, Professor of Gynecology and Obstetrics University St. Clement of Ohrid, Bitola, North Macedonia.

Received: February 21, 2022; **Published:** April 29, 2022

Abstract

PTBs are a significant global health issue worldwide and a leading cause of perinatal morbidity and mortality. Perinatal infections are the leading cause of PTBs. Genital tract infections account for about 25 - 40% of preterm deliveries. Chorioamnionitis (CA) is considered an important reason for prematurely childbirth, as microbiological data testify. Thus there are several diagnoses in the literature. Histopathological diagnosis CA defines as the presence of inflammatory cells in the fetal membranes. Clinically, CA is defined as Typically maternal fever and at least two of next signs: uterine tenderness, fetal and/or maternal tachycardia, maternal leukocytosis, foul smelling fluid. A number of bacterial, viral, fungal, and parasitic pathogens may be etiological factors of CA. In term births, CA is present in 4% of births. However, that frequency is higher in preterm births and PPRM. In pregnant women with clear clinical signs of CA, there is a strong correlation between histologically confirmed CA and the cardinal symptoms of CA. Often, the diagnosis of CA is delayed, leading to delayed preterm delivery, endangering the health of the pregnant woman, and damage to the fetus. Therefore, early recognition and diagnosis of CA is especially important. The diagnosis is made on the basis of clinical maternal symptoms. The primary management of CA is administration of antibiotics. Immediate treatment with antibiotics and proper care will ensure a better outcome of the CA-threatened pregnancy.

Keywords: *Preterm Birth; Chorioamnionitis; Diagnosis; Management*

Introduction

Preterm births (PTBs) are all births before 37 completed weeks or 259 days of gestation. It is a major determinant of neonatal mortality and morbidity and has long-term adverse consequences for health [1,2]. PTBs are a significant global health issue worldwide and a leading cause of perinatal morbidity and mortality. The incidence of PTB has not changed during the last 50 years. Every year about 15 million babies are born preterm. PTB usually affects 5 - 7% of births, but most of them are in developing countries [3]. The incidence of PTB has increased worldwide. In 2010, 11% of all live births were born preterm [4]. It is very important to identify women and pregnant at risk for PB early in pregnancy.

Genital tract infections account for about 25 - 40% of preterm deliveries. Women with Chlamydia trachomatis, Gardnerella Vaginalis, Trichomonas Vaginalis, Neisseria Gonorrhoeae, Treponema Pallidum, have a higher rate of preterm births. Still, the infection is difficult to detect due to the limitations of conventional microbial techniques and the difficulties in obtaining appropriate diagnostic samples during

pregnancy [6,7]. Experimental models also suggest the possible induction of labor by viral infection [5]. Chorioamnionitis is considered an important reason for prematurely childbirth, as microbiological data testify [5-7].

Definition

Chorioamnionitis (CA) is technically a histopathological diagnosis, but clinically the term “intraamniotic infection” may be more appropriate. Thus there are several diagnoses in the literature. Histopathological diagnosis CA defines as the presence of inflammatory cells in the fetal membranes. Clinically, CA is defined as Typically maternal fever and at least two of next signs: uterine tenderness, fetal and/or maternal tachycardia, maternal leukocytosis, foul smelling fluid. Taking into account the maternal inflammatory response, CA is defined as: Inflammatory changes limited to the subchorion, chorion and amnion. Finally, if the fetal immune response is considered, CA will be defined as umbilical vasculitis, funistis, elevated inflammatory markers in the cord blood. However, CA can be defined as infection and inflammation of the chorionic membrane (chorionic and amniotic) and/, or the amnion/amniotic fluid. CA is most commonly caused by a maternal ascending polymicrobial pathway, and can be infected by the fetus following exposure to an infection, directly by contact with amniotic fluid, or indirectly by placental-fetal circulation [8,9].

Etiology

A number of bacterial, viral, fungal, and parasitic pathogens may be etiological factors of CA. The most common causes are: *Ureaplasma urealyticum* (15 - 62%), *Mycoplasma hominis* (7 - 35%), Group B *Streptococcus* (8 - 11%), *Escherichia coli* (7 - 12%), *Gardnerella vaginalis* (8 - 25%), *Bacteroides sppeties* (8 - 30%), *Fusobacterium sppeties* (10 - 67%), *Prevotella sppeties* (17%), *Peptostreptococcus sppeties* (16%) etc [9].

CA is most often due to ascending bacteria causing acute inflammation of the membranes and the chorion of the placenta. Risk factors for CA include longer duration of membrane rupture (more than 12 h), nulliparity, internal monitoring of labor, multiple vaginal exams (more than or equal to 3), smoking, alcohol or drug abuse, immune compromised states and colonization with group B streptococcus. PPRM is the main factor of risk for clinical manifestation of CA. However, it should not be forgotten that preterm delivery and PPRM are most often a consequence of subclinical CA [10].

CA is an infection that can occur before labor, during labor. It can be acute, subacute or chronic. The effects of CA on the health of the newborn are particularly important in clinical practice. In the case of premature birth, the newborn is already at risk of all the risks of early termination of pregnancy. Additionally, CA carries additional health risks for the newborn. Most commonly, CA is associated with chronic lung disease in the infant, premature retinopathy, very low birth weight, and impaired brain development [10].

Epidemiology

In term births, CA is present in 4% of births. However, that frequency is higher in preterm births and PPRM. Premature delivery and PPRM, histologically in addition to CA, are also dominated by vasculitis [1-13]. Histologically confirmed CA is present in 94% of births that occurred between 21-24 weeks of gestation [14,15].

Pathogenesis

CA is most often an ascending infection, originating from the lower urogenital tract (cervical or vaginal infection). Other ways of transmitting the infection to the chorion and amnion have already been mentioned above.

A pathological feature of CA is neutrophil infiltration into fetal membranes and the presence of neutrophils in amniotic fluid.

Whether the infection is ascending or descending, the first step in the pathogenesis of CA is the passage of infectious agents into the chorioamnion and/or umbilical cord and/or placenta. The presence of infectious agents in the chorioamnion causes an inflammatory response in the mother and fetus, characterized by the release of a combination of proinflammatory and inhibitory cytokines and chemokines in both the mother and the fetus. The inflammatory response may cause clinical chorioamnionitis and/or lead to prostaglandin release, cervical maturation, membrane rupture and delivery (term or preterm delivery). In conditions of CA, in addition to the risk of direct adverse effect of the infectious agent on the fetus, one must not forget the effect of the fetal inflammatory response which may cause cerebral white matter damage, which damage may cause cerebral palsy, short-term or long-term neurological sequelae [10,11,13].

The consequences of maternal CA are manifested as: endomyometritis, wound infection, pelvic abscess, bacteremia and postpartum-hemorrhage, while, septic shock, disseminated intravascular coagulation, adult respiratory distress syndrome and maternal death are only rarely encountered [10,16,17].

Clinical signs and symptoms

CA is clinically manifested as a maternal fever associated with an elevated white blood cell count ($\geq 15,000/\text{mm}^3$), uterine tenderness (4 - 25% cases), abdominal pain, purulent fluid coming from the cervical canal, and fetal and maternal tachycardia (in 50 - 80% cases) and fetal tachycardia (in 40 - 70% cases). Presence of maternal fever is a necessary criterion for the diagnosis of clinical CA (maternal temperature $\geq 38^\circ\text{C}$). Fever is present in 95 - 100% of cases of clinical CA and is typically required for the diagnosis. The majority of women presenting with CA are in labor or have ruptured membranes [10,11,13,15].

Diagnosis

Often, the diagnosis of CA is delayed, leading to delayed preterm delivery, endangering the health of the pregnant woman, and damage to the fetus. Therefore, early recognition and diagnosis of CA is especially important.

The diagnosis is made on the basis of clinical maternal symptoms. Laboratory signs in favor of CA are: elevated leukocyte count, elevated CRP, microbiological examination of vaginal and cervical material. Postpartum, histopathological diagnosis is made by analysis of the chorion and amnion, placenta, and umbilical cord [10,11,14].

Differential-diagnostic, in women with fever, pain, and tenderness during labor must be evaluated for other common infections such as appendicitis, urinary tract infection, pyelonephritis, and pneumonia. Additional ancillary testing and examination must be thoroughly reviewed before making a final diagnosis [10].

Management

The primary management of CA is administration of antibiotics. The most common antibiotics used are ampicillin and gentamicin. Alternative antibiotics include clindamycin, metronidazole, cephalosporins, and vancomycin in women allergic to penicillin. After delivery, the current recommendation is to administer one additional dose with a cesarean section but no additional antibiotics for vaginal deliveries. Additional broad-spectrum antibiotics may be required, depending on the clinical status [10,18].

Conclusion

CA is a common and serious perinatal problem. If not recognized in time and treated appropriately and in a timely manner, CA leads to significant maternal, fetal, and neonatal morbidity and mortality. The diagnosis is based on the clinical signs, and therefore it is important during the perinatal examinations, to educate the pregnant woman what are the symptoms that if they occur, she should immediately

contact a gynecologist. Immediate treatment with antibiotics and proper care will ensure a better outcome of the CA-threatened pregnancy. Research in this area and the discovery of new biomarkers will allow appropriate interventions and treatments to be applied in the future, thus contributing to the prevention and more effective treatment of CA.

Bibliography

1. Goldenberg RL, et al. "Epidemiology and causes of preterm birth". *Lancet* 371 (2008): 75-84.
2. Wang ML, et al. "Clinical outcomes of near-term infants". *Pediatrics* 114 (2004): 372-376.
3. Lawn JE, et al. "One year after The Lancet Neonatal Survival Series - was the call for action heard?" *Lancet* 367 (2006): 1541-1547.
4. Blencowe H, et al. "National, regional, and worldwide estimates of preterm birth rates in the year 2010 with time trends since 1990 for selected countries: a systematic analysis and implications". *Lancet* 379 (2012): 2162-2172.
5. Jasovic-Siveska E. "Editorial: Etiology, Prevention and Prediction of Preterm Birth- What is New?" *Obstetrics and Gynecology International Journal* 8.3 (2017): 00291.
6. Jasovic-Siveska E. "Prevention and Prediction of Preterm Birth-Status Quo in the Last 50 Years". *Reproductive System and Sexual Disorders* 3 (2014): e117.
7. Lamont RF. "Infection in the prediction and antibiotics in the prevention of spontaneous preterm labor and preterm birth". *BJOG: An International Journal of Obstetrics and Gynaecology* 110.20 (2003): 71-75.
8. Stojanovska V, et al. "The Consequences of Preterm Birth and Chorioamnionitis on Brainstem Respiratory Centers: Implications for Neurochemical Development and Altered Functions by Inflammation and Prostaglandins". *Frontiers in Cellular Neuroscience* 12 (2018): 26.
9. Ericson JE and Laughon MM. "Chorioamnionitis: implications for the neonate". *Clinics in Perinatology* 42.1 (2015): 155.
10. Tita AT and Andrews WW. "Diagnosis and management of clinical chorioamnionitis". *Clinics in Perinatology* 37.2 (2010): 339-354.
11. Suzuki S. "Association between clinical chorioamnionitis and histological funisitis at term". *JNPM Journal of Neonatal-Perinatal Medicine* 12.1 (2019): 37-40.
12. Kim B, et al. "Placental Lesions in Meconium Aspiration Syndrome". *Journal of Pathology and Translational Medicine* 51.5 (2017): 488-498.
13. Kim CJ, et al. "Acute chorioamnionitis and funisitis: definition, pathologic features, and clinical significance". *American Journal of Obstetrics and Gynecology* 213.4 (2015): S29-52.
14. Perkins RP, et al. "Histologic chorioamnionitis in pregnancies of various gestational ages: implications in preterm rupture of membranes". *Obstetrics and Gynecology* 70.6 (1987): 856-860.
15. Conti N, et al. "Term histologic chorioamnionitis: a heterogeneous condition". *European Journal of Obstetrics and Gynecology and Reproductive Biology* 188 (2015): 34-38.
16. Saghafi N, et al. "Cervical bacterial colonization in women with preterm premature rupture of membrane and pregnancy outcomes: A cohort study". *International Journal of Reproductive BioMedicine* 16.5 (2018): 341-348.

17. Shanks AL, *et al.* "Treatment Utility of Postpartum Antibiotics in Chorioamnionitis Study". *The American Journal of Perinatology* 33.8 (2016): 732-737.
18. Committee Opinion No. 712. "Intrapartum Management of Intraamniotic Infection". *Obstetrics and Gynecology* 130.2 (2017): e95-e101.
19. Kemp MW. "Preterm Birth, Intrauterine Infection, and Fetal Inflammation". *Frontiers in Immunology* 5 (2014): 574.

Volume 11 Issue 5 May 2022

©All rights reserved by Emilija Jasovic-Siveska.