Brucella Infection Causing Abortion in Human Beings

E. Zowghi, A. H. Hedayeti¹, A. Ebadi, A. M. Behroozikhah and M. Yarahmadi

Razi Vaccine and Serum Research Institute, PO Box 11365-1558, Tehran, Iran.
1- Feiaz-Bakhsh Hospital, Karadj, Iran

Summary

In a survey on human brucellosis, in a period of 2 years, natural abortions were investigated for Brucella. A total of 67 foetuses and 203 placentas were collected from Feiaz-Bakhsh Hospital, Karadj. These samples were bactriologically examined and Brucella was isolated from 1 foetus and 1 placenta. Both isolates were classified by biovar procedures and were found to be B. melitensis biovar 1. The mother of the positive foetus, when serolgically tested for antibodies to Brucella, showed a high titre. She had had 2 abortions during previous 18 months. It was postulated that Brucella was, possibly, the cause of abortion in the pregnant women.

Introduction

Localisation of *Brucella* micro-organism in genital organs of animals, particularly ruminants, is a well known fact. In these species, the most intensive lesions occur in the tissues of pregnant uterus and, usually, the result is abortion in the second third of pregnancy. The presence of erythritol in the placenta of animals has been determined as the *Brucella* growth stimulant resulting in abortion due to the multiplication of *Brucella* and the subsequent necrosis and destruction of foetus membranes (Nielson and Duncan, 1990). Although erythritol is absent in the human placenta, the occurrence of abortion due to brucellosis in pregnant women, particularly in the second third of pregnancy, still is a possibility. Likely, the abortion occurs following acute fever and bacteraemia of the mother and toxaemia of the foetus.

Despite the fact that *Brucella* can survive and multiply within the host cells, large number of them undergo destruction and endotoxin is released. In addition to the effect of endotoxin on various tissues of the body, abortion

directly due to endotoxin occurs in animals and, occasionally, in human beings. Therefore, many workers believe that in premature parturition transmission of *Brucella* from placenta to the infant is a possibility (Madkur, 1989). Brucellosis in human beings occurs with involvement of various organs and a variety of manifestations. Infection of the genitourinary system is predominant in animals but accounts for less than 20% of complications of brucellosis in the human. The *Brucella* organisms seem to have a predilection for the reproductive organs of both sexes in persons and animals. They occasionally localise in the human female genital tract. Tubovarian and pelvic abscesses, chronic salpingitis, cervicitis and menstrual disturbances are reported (Spink, 1956; Porreco and Haverkamp, 1974).

In the present study, the results of bacteriological examination of 67 foetuses and 203 placentas are presented.

Materials and methods

In a period of 2 years, from the beginning of 1993 to the end of 1994, sixty-seven foetuses and 203 placentas were collected from Feiaz-Bakhsh Hospital, Karadj, and studied at Razi Vaccine and Serum Research Institute. Samples of spleen, liver, lungs and stomach contents of all foetuses and 4-5 cotyledons of each placenta were cultured on agar plates of serum dextrose medium with antibiotics. All plates were incubated at 37°C in a carbon dioxide incubator for *B. abortus*, and in ordinary incubator for *B. melitensis*. They were examined 4 to 7 days later for *Brucella*-like colonies. The isolates were tested for smooth and rough colonies and examined for agglutinablity by using *B. abortus* and *B. melitensis* monospecific antiserum. Subcultures of colonies were made on *Brucella* agar slopes and incubated at 37°C. The isolates, after being tested for purity, were biotyped using techniques recommended by Alton *et al.* (1988) and Young and Corbel (1989). The sera of some patients were examined for brucellosis by serological tests.

Results

Of the 67 foetuses tested during the period of the study, 1 case was positive for *Brucella* organism. Also, out of 203 placenta-samples examined in this period, 1 case was positive for *Brucella*. Both isolates were identified, by sensitivity to dyes, failure of H₂S production, agglutination with *B. melitensis* monospecific antiserum and lack of Tb phage sensitivity. The mother of the positive foetus had a titre of 1:320 in serum agglutination test

and a titre of 1:80 in 2-Mercaptoethanol test. She had had 2 other abortions during the previous 18 months.

Discussion

The role of bacterium Brucella in causing human abortions has not been evaluated by appropriate case-control studies, therefore, it is not yet known whether brucellosis causes abortions in pregnant women more frequently than other bacterial infections (Spink, 1956). Although human placenta does not contain erythritol, the fact that Brucella has been recovered from placental tissues and amniotic fluid in women with acute brucellosis indicates that there may be some risk of abortion during the course of brucellosis. The first report on the Brucella infection of human uterus dates back to 1907 (Williams, 1907). Perusal of the literature up to 1980s show that Brucella organisms have been isolated from human foetuses on numerous occasions (De Forest, 1917; Kristensen, 1929; Carpenter and Boak, 1931; De Carle, 1931; Janbon and Kerleau, 1939; Hagebusch and Frei. 1941; Williamson, 1944; Poole et al., 1972; Sarram et al., 1974; Schreywer et al., 1980). Oran et al. (1983) reported a premature baby, with B. abortus infection, whose mother had a high Brucella agglutination titre and had given birth to another premature baby the year before. In 1985, many cases of abortions due to B. melitensis were reported from Saudi Arabia (Mohammed et al., 1985; Madkur et al., 1985). Lubani et al. (1988) reported three Arab new-borns whose mothers had Brucella infection during pregnancy. Other cases of abortions and premature new-borns were reported in 1990 and 1992 (Labrune et al., 1990; Al-Eissa and Mofada, 1992). The difference in manifestation of brucellosis in ruminants and the man, that ruminants but not human beings routinely abort their foetuses, is attributed to the presence of erythritol in the ruminant placenta. However, transmission of infection through placenta to the foetus and isolation of Brucella from foetus or premature new-born and placenta have been frequently reported in human beings (Al-Eissa and Al-Zamil, 1991; Al-Eissa and Mofada, 1992; Benjamin and Annobil, 1992) and is prevalent in endemic areas. The isolation of B. melitensis from 2 abortion cases corroborates the findings by other workers. It is inferred that Brucella can infect human chorioamniotic tissues during pregnancy and may induce abortion. However, this is not unique to Brucella and generally reflects the severity of bacterial infection, it may occur following infection by many other micro-organisms.

Acknowledgements

The co-operation of the staff of Feiaz-Bakhsh Hospital, particularly Mrss. Chavoshi, Mechaniki, Dehghan-Shoar, Nassajan, Farahnood and Sadeghian are gratefully acknowledged. Also we are very grateful to Mrss. Z. Bashir-Hashemi and Kh. Kobari and the rest of the staff at Brucellosis Department in Razi Vaccine and Serum Research Institute for their technical assistance.

References

- Alton, G.G., Jones, L.M., Angus, R.D. and Verger, J.M. (1988). Techniques for the brucellosis laboratory. IRAA. Paris
- Benjamin, B. and Annobil, S.H. (1992). Childhood brucellosis in southwestern Saudi Arabia: A 5 year exprience. Journal of Tropical Pediatrics, 38: 167-172
- Carpenter, C.M. and Boak R. (1931). Isolation of *Brucella abortus* from a human foetus. Journal of Ameriacan Medical Association, 96: 1212-1216
- De Carle, D.W. (1931). Premature labour in the case of undulant fever. Mayo Clinical proceedings, B 6: 469-471
- De Forest, H.P. (1917). Infectious abortions of cattle as a complication or pregnancy for women. American Journal of Obstetrics, 76: 221-229
- Hagebusch, O. and Frei, C.F. (1941). Undulant fever in children. American Journal of Clinical Pathology, 11: 497-515
- Janbon, M. and De Kerleau, J.C. (1939). Brucellose humaine et avortement: Isolment de *Brucella melitensis* dans le sang de la mer, les visceras, et le sang du foetus, le placenta, et les lochies. Presse Medical, 47: 453-457
- Klaus, N. and Duncan, J.R. (1990). Animal Brucellosis. New York, CRC press
- Kristensen, M (1929). La position bacteriologique du Bacillus *abortus* de Bang, son importance comme cause de maladie chez l'homme. Annal Medicine, 26: 339-350
- Labrune, P., Jabir, B., Magny, J.F., Guilbert, M., Damay, M. and Odievre, M. (1990). Recurrent enterocolitis-like symptoms as the possible presenting manifestations of neonatal *Brucella melitensis* infection. Acta paediatric. Scandinavica, **79**: 707-709
- Lubani, M.M., Dudin, K.L., Shrada, D.C., Sinna, N.M. (1988) Neonatal brucellosis. European Journal of Pediatrics, 147: 320-522
- Madkur, M. M. (1989). Brucellosis. London, Butterworths press
- Madkur, M.M., Mohamed, A.E., Talukder, M.A.S. and Kudwah, A. (1985) Brucellosis in Saudi Arabia. Saudi Medical Journal, 6: 324-332
- Mohamed, A.E., Madkur, M.M., Talukder M.A.S. and Hifzi, I. (1985). Obstetrical and gynaecological presentations in brucellosis. Arab Journal of Medicine, 4: 22-26
- Oran, O., Kaslioglu, A., Seemeer, G. (1983). A premature infant with *Brucella abortus* infection. Turkish Journal of Paediatrics, 25: 139-142
- Poole, P.N., Whitehouse, D.B. and Gilchrist, M.M. (1972). A case of abortion consequent upon infection with *Brucella abortus* type 2. Journal of Clinical pathology, 25: 882-884

- Porreco, R.P. and Haverkamp, A.D. (1974). Brucellosis in pregnancy. Obstetrics and. Gynecology, 44: 597-602
- Sarram, M., Feiz, J., Foruzandeh, M. and Gazanfarpour, P. (1974). Intrauterine foetal infection with *Brucella melitensis* as a possible cause of second-trimester abortion. American Journal of Obstetrics and Gynecology, 119: 657-660
- Schreyer, P., Caspie, L. Y, Eshchar, Y. and Sompolinsky, D. (1980). *Brucella* septicaemia in pregnancy. European Journal of Obstetrics, Gynecology and Reproduction, 10: 99-107
- Spink, W.W. (1956). The nature of brucellosis. Minneapolis, MN: University of Minnesota press
- Williams, E.M. (1907). Mediterranian fever: Infection in utero. Journal of Royal Army Medical Corps, 9: 59-60
- Williamson, A.R.H. (1944). Abortions in *Brucella abortus* fever. British Medical Journal, 1: 752-756
- Young, E.J., and Corbel, M.J. (1989). Brucellosis: clinical and laboratory aspects. New York, CRC Press
- Yousef Al-Eissa, Fahad Al-Zamil, Mohamed Al-Mugeiren (1991). Childhood Brucellosis: A deceptive infectious disease. Scandinavian Journal of Infectious Diseases, 23: 129-133
- Youssef, Al-Eissa and Saleh, M.M. (1992). Congenital Brucellosis. Pediatrics Infectious Disease Journal, 11: 667-671