

## 1 **AMNIOINFUSION: FROM TERMINATION OF PREGNANCY TO THERAPY**

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3 Amnioinfusion or infusion of saline into the amniotic cavity was first used as a  
4 technique for late termination of pregnancy in the 1960s (Jonas and Slate. *Obstet*  
5 *Gynecol.*1966;27:494-6). By the mid 1970s, transabdominal amnioinfusion of  
6 hypertonic (20%) saline was the most widely used method of pregnancy termination  
7 after 14 weeks in the US. This technique was associated with around 10% morbidity  
8 including hemorrhage requiring transfusion, retained tissue requiring surgical removal,  
9 infection, coagulopathy, and hypernatremia (Ballard and Ballard. *Am J Obstet*  
10 *Gynecol.*1972;114:575-81). There were also risks of intravenous, intraperitoneal, or  
11 intramyometrial injection of the saline and thus amnioinfusion for abortion was  
12 abandoned in the 1980s in favour of prostaglandins (PGE2) which were more efficient  
13 and with fewer side effects.

14 Miyazaki and Taylor were the first to report on the use of saline amnioinfusion  
15 in 42 patients having repetitive variable or prolonged decelerations that did not  
16 respond to conventional therapy, such as maternal position changes and oxygen  
17 administration (Miyazaki and Taylor. *Am J Obstet Gynecol.*1983;146:670-8). The  
18 technique used normal saline or ringers lactate infused transcervically through a  
19 catheter into the uterine cavity, or transabdominally using a 'spinal' needle when  
20 membranes are intact. More recently, transcervical amnioinfusion was proposed for  
21 women presenting with thick meconium staining of the amniotic fluid to reduce the risk  
22 of perinatal death and meconium aspiration syndrome. Although the first results  
23 seemed encouraging (Hofmeyr et al.*BJOG.*1998;105:304-8), the 2006 UK National  
24 Institute for Health and Care Excellence (NICE) concluded that there was insufficient  
25 evidence to support the practice ([www.nice.org.uk](http://www.nice.org.uk) › guidance › ipg192) and it has  
26 rarely been used for this indication. More recent data from India have suggested that

27 transcervical amnioinfusion in labour for meconium-stained amniotic fluid can be  
28 performed safely in a setup with limited neonatal care facilities, to decrease the  
29 incidence of caesarean deliveries and improve neonatal outcome (Choudhary and  
30 Bano. Arch Gynecol Obstet.2010;282:17-22).

31 Amnioinfusion was also used prophylactically in various conditions commonly  
32 associated with oligohydramnios to limit the risk of cord compression or  
33 oligohydramnios-related pulmonary hypoplasia (Figure) (Fisk et al.  
34 BJOG.1992;99:464-8) but was not found to improve neonatal outcomes. Two large  
35 trials have assessed whether infusion of fluid into the amniotic cavity could improve  
36 pregnancy outcome after second-trimester premature rupture of the membranes  
37 (AMIPROM trial Roberts et al.UOG 2014;43:490-9;PPROMEXIL-III trial van Teeffelen  
38 et al. BMC Pregnancy Childbirth. 2014; Apr 4;14:128). The first trial suggested an  
39 improvement in long-term healthy survival after amnioinfusion but neither of these  
40 studies showed a significant better outcome after amnioinfusion. In this issue, De  
41 Ruigh et al. (BJOG 2021) report on the long-term outcome of the PPRMEXIL-III trial.  
42 Overall, of the 14 survivors (all born at a median gestational age of 24 weeks), 71%  
43 had no neurodevelopmental delay. The small sample size of this study does not allow  
44 to draw any definitive conclusions and survival without developmental delay nor  
45 respiratory problems did not differ between the treatment arm and controls with no  
46 amnioinfusion. From termination of pregnancy to therapy, amnioinfusion has still to  
47 find its place in modern obstetrics.

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49 Word count: 497

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52 BJOG since 1902 Perspectives on BJOG-19-0283R2

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55 **Disclosure of interests**

56 The authors declare no conflicts of interest.

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58 Elena Contro, University Hospital of Bologna, Italy.

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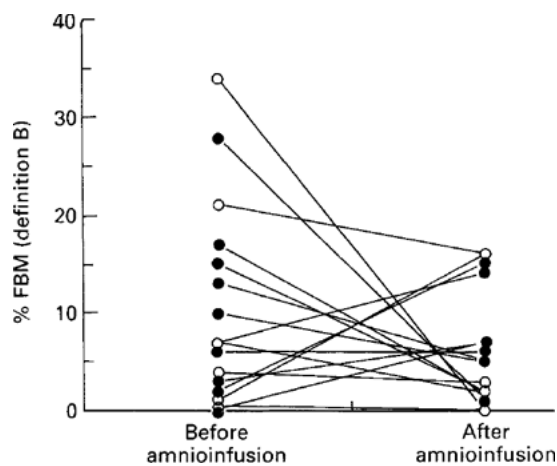
60 Eric Jauniaux, Professor, EGA Institute for Women's Health, Faculty of Population

61 Health Sciences, University College London, UK.

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63 Both authors are BJOG scientific editors

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68 **Figure:** The incidence of FBM (= % of time spent in FBM epochs) using definition B

69 before and after amnioinfusion in 16 pregnancies. ○ pregnancies that leaked; ●

70 pregnancies that retained infused fluid (From Fisk et al., BJOG.1992;99:464-8).

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