

**Appendix 6 (as supplied by the authors): Risk of spontaneous abortion associated with the use of antibiotics during pregnancy, by drug classes from the beginning of gestation until 15 days before index date<sup>\*\*</sup>.**

| Exposure   | no (%) of cases | Crude OR<br>(95% CI) | Adjusted OR<br>(95% CI) |
|--|-----------------|----------------------|-------------------------|
| Class of antibiotics                               |                 |                      |                         |
| No use   | 7470 (85.84%)   | 1.00                 | 1.00                    |
| Cephalosporins                                     | 58 (0.67%)      | 1.11(0.85-1.45)      | 1.03(0.79-1.35)         |
| Penicillins  | 429 (4.93%)     | 0.94(0.86-1.03)      | 0.86 (0.77-0.96)        |
| Macrolides   | 231(2.65%)      | 1.73(1.50-1.98)      | 1.56(1.35-1.80)         |
| Quinolones   | 160(1.84%)      | 3.01(2.53-3.59)      | 2.72(2.26-3.27)         |
| Sulfonamides                                       | 29 (0.33%)      | 2.27(1.53-3.37)      | 2.12(1.42-3.17)         |
| Tetracyclines                                      | 55(0.63%)       | 2.18 (1.63-2.90)     | 2.05 (1.52-2.75)        |
| Other antibacterials                               | 31(0.36%)       | 1.34 (0.94-1.93)     | 1.18 (0.78-1.78)        |
| Antiprotozoals                                     | 49(0.56%)       | 2.00(1.48-2.71)      | 1.85(1.36-2.50)         |
| Urinary anti-infectives                            | 25 (0.29%)      | 0.58(0.39-0.86)      | 0.56 (0.38-0.82)        |
| Combined use of $\geq 2$<br>classes of antibiotics | 165 (1.90%)     | 1.67(1.42-1.96)      | 1.44(1.22-1.71)         |

Note: 95%CI, 95% confidence interval; OR, odds ratio.

<sup>\*\*</sup>Adjusted for the following variables: maternal age on the first day of gestation, maternal marital status (living alone or cohabiting), receipt of social assistance during pregnancy, education level in years ( $\leq 12$  or  $>12$ ), and area of residence on the 1DG (urban or rural); maternal chronic co-morbidities assessed using physician-based diagnoses or filled prescriptions of related medications in the year before and during pregnancy until index date (chronic hypertension, depression, diabetes mellitus, asthma, epilepsy, polyarthritis rheumatoid and systemic lupus erythematosus, thyroid disorders); Uterine malformations and endometriosis assessed using physician-based diagnoses; Maternal infections assessed using physician-based diagnoses in the year before and during pregnancy until index date (urinary tract infection, respiratory tract infection, bacterial vaginosis, and sexually transmitted diseases); Other anti-infective agents assessed also in the year before and during pregnancy until index date; Use of health services in the year before pregnancy; and history of planned and spontaneous abortion.