

Trends of influenza vaccination coverage in pregnant women, a ten-year analysis from a French healthcare database

Mélodie Corbeau

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UNIVERSITÉ CLERMONT AUVERGNE UFR DE MÉDECINE ET DES PROFESSIONS PARAMÉDICALES

THÈSE D'EXERCICE

pour le

DIPLÔME D'ÉTAT DE DOCTEUR EN MÉDECINE

par

CORBEAU, Mélodie

Présentée et soutenue publiquement le 22 juin 2021

Trends of influenza vaccination coverage in pregnant women, a ten-year analysis from a French healthcare database

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Dédicaces validées le : & /o5/202

Signature du directeur de thèse :

Mélodie & ORBEAU

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Liste des abréviations

ALD: Affection Longue Durée

ATC: Anatomical Therapeutic Chemical class

CCAM: Common Classification of Medical Acts

CI: Confidence Interval

CMUc : Couverture Maladie Universelle complémentaire

CNIL: Commission Nationale Informatique et Libertés

COVID-19: Coronavirus Disease 2019

C-section: Cesarean-section

DTP: Diphtérie Tétanos Polio

ECDC: European Centre for Disease Prevention and Control

EPICES : Evaluation de la Précarité et des Inégalités de santé dans les Centres d'Examens de

Santé

EGB: Echantillon Généraliste des Bénéficiaires

HIV: Human Immunodeficiency Virus

ICD-10: International Classification of Diseases, Tenth revision

IVC: Influenza Vaccination Coverage

OR: Odd Ratio

SPF: Santé Publique France

T1,2, 3: first, second or third Trimester of Pregnancy

UK: United Kingdom

USA: United States of America

WHO: World Health Organization

1 Introduction

Influenza is an acute respiratory infection caused by *Myxovirus influenza*, a virus whose subtypes A and B are responsible for seasonal outbreaks worldwide. The World Health Organization (WHO) estimates at a billon the number of influenza cases every year, 3 to 5 million severe cases and 290 000 to 650 000 deaths [1].

Pregnant women are more likely to develop a severe form of influenza [2–6] associated with obstetrical complications such as prematurity and miscarriage [7,8].

Every year, a vaccine is developed with the latest epidemiological information on the circulating subtype in the southern hemisphere. The efficacy and innocuity of the vaccine during pregnancy have been well established in numerous studies [9–20], as well as its protective effect for the infant through passive immunization [21–23].

In Europe the majority of countries have recommended vaccination for pregnant women, whatever the term of pregnancy [24], since the WHO issued its recommendation in 2012 [25]. In France, the influenza vaccination has been recommended for pregnant women since 2009 (in the 2nd and 3rd trimester) and generalized in 2012 to all stages of pregnancy [25,26]. Therefore, it should be prescribed to all pregnant women, to avoid severe forms of influenza and to protect their infant.

However, a report issued by the European Centre for Disease Prevention and Control (ECDC) in 2017 showed that the influenza vaccination coverage (IVC) in pregnant women in Europe remained low and uneven, from 0.3% to 56.1% in 2014-2015 and was not monitored by most of the member states, including France [24]. A few studies all based on classical surveys with different methodologies estimated the IVC in France to be between 5.4% and 26% [27–30]. In order to avoid the usual biases arising from classical surveys (selection bias and generalization

of the findings), leading to a great variability in coverage estimates, and since the influenza vaccination is prescribed and fully reimbursed by the social security, we propose to use the French national healthcare insurance database to provide estimates of the IVC in pregnant women.

In this study we sought to analyze the evolution of the annual IVC in pregnant women in France from 2009 to 2018, to evaluate the impact of the WHO 2012 recommendation [25] on vaccination behavior, and to identify determinants of influenza vaccination uptake in pregnancy.

2 Methods

2.1 Data source and design

A repeated annual cross-sectional study was performed by using data from the "Echantillon Généraliste des Bénéficiaires" (EGB), a representative sample of the French population, from January 1, 2009 to December 31, 2018. The EGB database contains the healthcare data of about 660 000 people, representing 1/97th of the French population. This sample of insured patients is randomized, anonymized and representative of the health care protected French population [31,32]. EGB contains sociodemographic information on the beneficiaries, as well as data on various health expenses reimbursed by the French healthcare insurance. That includes prescribed medications (identified according to their anatomical therapeutic chemical class, ATC), consultation with medical and para-medical professional, in private practice as well as public hospital, hospitalization and laboratory expenses. We can also find data on chronic diseases such as ALD (long-term disease) codes, diagnosis and medical history coded during hospitalization using ICD-10 (international classification of diseases, tenth revision) and medical procedures referenced according to the CCAM (common classification of medical acts). The data have been collected since 2005 and are available after a two-year period.

The EGB database is a validated tool for pharmacoepidemiological studies in France and has already proven to be a reliable tool to assess vaccination coverage for other reimbursed vaccines [33–35].

2.2 Study population

All women who had data recorded on a delivery, vaginal birth, or caesarian section in the EGB database, during the period from January 1, 2009 to December 31, 2018 were included. The exclusion criteria were unwanted pregnancy and denial of pregnancy. We did not exclude pregnancies that resulted in stillbirth, or therapeutic abortion after 22 weeks of gestation.

The corresponding ICD 10 diagnosis codes and CCAM medical procedures are listed in supplementary tables 1-3.

2.3 Exposure

Influenza vaccine exposure of the pregnant women included was identified through the reimbursement for a seasonal influenza vaccine in the EGB database, up to nine months before the date of their delivery, using two codes indicating dispensation (ATC J07BB and PRS_NAT 3331). We assumed that a vaccine purchased was injected afterwards and we calculated the IVC based on the number of pregnant women who had a reimbursed influenza vaccine.

To evaluate the impact of the WHO 2012 recommendation [25] on vaccination in the first trimester, we formed 3 subgroups of women according to their term of pregnancy during the vaccination campaign that takes place from October to January. Women who gave birth in January were labeled T3 as they were in their third trimester during the vaccination campaign. Likewise, those who gave birth in April were labelled T2, and those who gave birth in July were labeled T1.

2.4 Collected data

The collected data concerned were pre-existing medical condition, age, parity, CMUc (universal complementary healthcare insurance), and prescriber's specialty.

We used the French influenza vaccination recommendation to select women with pre-existing conditions who would have been targeted by the vaccination campaign outside of their pregnancy. Various conditions were included, and grouped in pulmonary, cardiac, neurological, renal, and hepatic disease, diabetes, and immunodeficiency (including HIV, stem cell and all organ transplantations, inflammatory and autoimmune diseases, hematological disease such as sickle cell disease and cancer).

Data were extracted from the EGB database using the CCAM, ICD-10, or long-term chronic disease codes (supplementary tables 5-7).

CMUc was used as a proxy for precarity since it is an insurance available only to patients with low income, to insure their access to healthcare free of charge. The women affiliated with the CMUc were considered socioeconomically deprived.

2.5 Data analysis

The population was described by means and standard deviations for continuous data and by numbers and percentages for categorical data.

Analysis of influenza vaccination of pregnant women was carried out using Chi squared test (or Fisher's exact test when appropriate) for categorical data and using Student's t-test for continuous data. Odds ratio (of vaccination) between groups are presented with their 95% confidence interval.

To validate the reliability of our data and the computation method of IVC, we compared the IVC calculated through EGB in patients over 65 years old to French official influenza vaccination data (Santé Publique France, SPF).

All tests were two-sided. A p-value < 5% was considered statistically significant, but the interpretations were based-on size of differences with clinical view (rather than just regarding the p-value), as significant tests do not always reflect clinically relevant differences when involving large samples. All analyses were performed using SAS Enterprise Guide (SAS Institute Inc, Cary, NC, USA).

2.6 Ethics approval

There was no requirement for ethical approval for this study. The EGB database guarantees the confidentiality and anonymity of all data (agreement by the French Data Protection Authority, CNIL, June 14, 2005).

3 Results

3.1 Validation data

Compared to the nation-wide official influenza vaccination data (SPF) (Figure 1), data from the EGB database showed a similar trend of influenza vaccination coverage in the French population over 65 years old.



Figure 1: Influenza vaccination coverage in the French population over 65 years old in EGB and French Public Health data (SPF).

EGB: Echantillon généraliste des bénéficiaires ; SP: Santé Publique France

3.2 Influenza vaccination coverage in pregnant women

From January 1, 2009 to December 31, 2018 a total of 73 314 pregnant women were identified in the EGB database. Fifty-two denial of pregnancy and 1 055 unwanted pregnancies were

excluded from our cohort. A total of 72 207 pregnant women were included in the study (Figure 2), whose characteristics appear in the Table I.

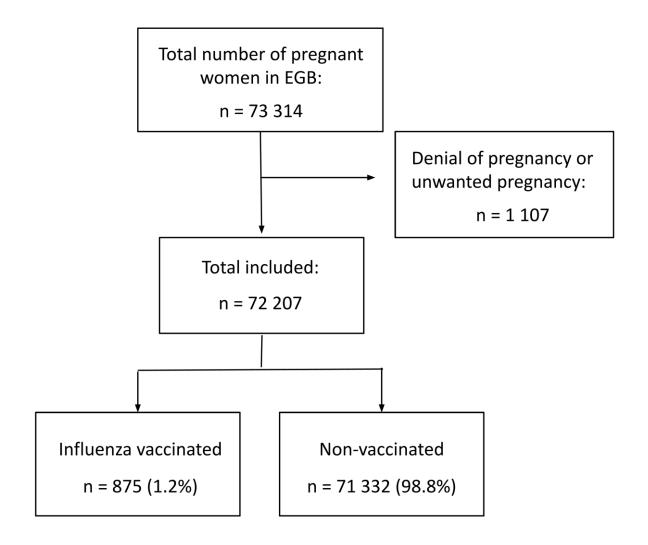


Figure 2: Flow chart of the study

The overall influenza vaccination coverage (IVC) was 1.21% in the 2009-2018 period (n=875/72 207; 95%CI 1.14-1.30). The IVC significantly increased after the 2012 recommendation, from 0.33% in the 2009-2012 period, to 1.79% in the 2013-2018 period (p<0.001) (Figure 3).

	Unvaccinated	Vaccinated			
	n (%)	n (%)	p-value	OR	95% CI
Total	71332 (100)	875 (100)			
Age	30.3 ± 5.40	32.0 ± 5.15	<0.001		
< 25	10162 (14.3)	63 (7.2)			
25-29	21691 (30.4)	215 (24.6)		1.60	1.21-2.12
30-34	23402 (32.8)	316 (36.1)		2.18	1.66-2.86
≥35	16075 (22.5)	281 (32.1)		2.82	2.14-3.71
Socioeconomic deprivation (CMUc)	12512 (17.5)	129 (14.7)	0.030	0.81	0.67-0.98
Preexisting condition	5847 (8.2)	150 (17.1)	<0.001	2.32	1.94-2.77
Pulmonary disease	1198 (1.7)	46 (5.3)	< 0.001	3.25	2.4-4.4
Cardiac disease	1633 (2.3)	45 (5.1)	< 0.001	2.31	1.7-3.13
Neurological disease	163 (0.2)	3 (0.3)	0.487	1.5	0.48-4.71
Renal disease	137 (0.2)	12 (1.4)	< 0.001	7.23	3.99-13.09
Sickle-cell disease	128 (0.2)	6 (0.7)	< 0.001	3.84	1.69-8.73
Diabetes	392 (0.5)	36 (4.1)	< 0.001	7.77	5.49-11
Immunodeficiency	1057 (1.5)	35 (4)	< 0.001	2.77	1.96-3.91
Hepatic disease	293 (0.4)	9 (1.0)	0.005	2.52	1.29-4.91
Obesity	2561 (3.6)	39 (4.5)	0.171	1.25	0.9-1.73
Multifetal pregnancy	1394 (2.0)	21 (2.4)	0.344	1.23	0.8-1.9
Parity			0.821		
1	34698 (48.6)	429 (49.0)		1.02	0.89-1.17
≥2	36634 (51.4)	446 (51.0)		0.98	0.86-1.12
Prematurity	6307 (8.8)	70 (8.0)	0.383	0.9	0.7-1.15
Still birth	203 (0.3)	1 (0.1)	0.345	0.4	0.06-2.86
C-section	14534 (20.4)	197 (22.5)	0.119	1.32	1.13-1.55
Therapeutic abortions	214 (0.3)	1 (0.1)	0.316	0.38	0.05-2.71

Table I: Influenza vaccination and associated factors among pregnant women from 2009 to 2018

OR: odd ratio calculated with Wald test

CI: confidence interval

CMUc: universal complementary healthcare insurance

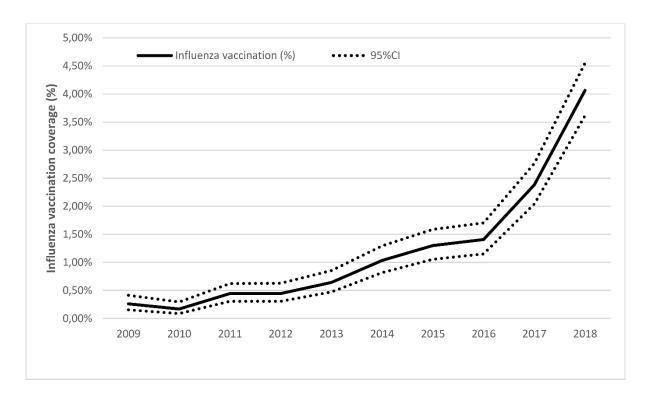


Figure 3: Evolution of the influenza vaccination coverage from 2009 to 2018

Vaccinated women were slightly older, less socially deprived, and had more pre-existing medical condition. There was no difference regarding the pregnancy outcome in the vaccinated and non-vaccinated population.

The mean gestational age at delivery was 38.9 weeks in both vaccinated and non-vaccinated populations. The mean gestational age of vaccination was 22.9 weeks \pm 9.7.

The identified prescribers were mainly family physicians (58%), followed by gynecologists (21%) and midwives (4%). Some specialists sporadically prescribed the vaccine (total of 5%) and 12% were dispensed directly by the pharmacists.

Women were most likely to be vaccinated if they were in their 2nd or 3rd trimester of pregnancy during the official vaccination campaign (Figure 4), but the vaccination rate increased for all three trimesters after the 2012 recommendation was issued.

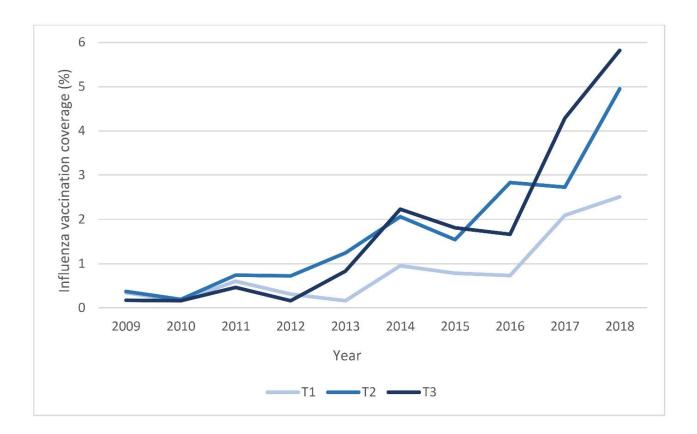


Figure 4: Evolution of influenza vaccination coverage by trimester of pregnancy during the vaccination campaign.

3.3 Influenza vaccination determinants

The vaccinated population was older than the non-vaccinated one (respectively 32 and 30.3 years old, p<0.001) and the vaccination rate increased proportionally with the age of the population (see Table I). Women younger than 25 years old had a low coverage (0.6%) whereas women over 35 years old were more likely to get the influenza vaccine (1.7%; OR: 2.82, 95%CI 2.14-3.71). Socio-economically deprived women were less likely to be vaccinated

(OR: 0.81, 95%CI: 0.67-0.98). Women with pre-existing medical conditions at risk for severe influenza had an overall higher vaccination rate (2.5%; OR: 2.32, 95%CI: 1.94-2.77). Of all the disease, renal disease and diabetes were associated with the highest vaccination rates (respectively 8.1% and 8.4%), and obesity with the lowest (1.5%).

The vaccination behavior was not influenced by multifetal pregnancy (OR: 1.23, 95%CI 0.8-1.9) or by parity: primiparous and multiparous women had the same vaccination rate (1.2%).

4 Discussion

4.1 Main results

To our knowledge, this is the first study to evaluate the trend of influenza vaccination coverage in pregnant women over such a long period (10 years) in a nationally representative population-based sample. Additionally, due to the large sample size, several correlates of influenza vaccination in pregnant women could be investigated.

Our primary endpoint was to analyze the evolution of the IVC in pregnant women in France from 2009 to 2018 using the French national healthcare database. We showed that the IVC increased significantly since the 2012 recommendation but remained very low (up to 4.1% in 2018). In addition, the coverage for women who started their pregnancy during the vaccination campaign remained lower than those of women in their 2nd and 3rd trimester, thus questioning the impact of the recommendation, that encouraged the vaccination of women at all term of pregnancy. The IVC was higher for older women, and women with pre-existing condition at risk for severe influenza, and lower in socially deprived women as often described in other studies [30,36–38].

The vaccines were mainly prescribed by family physicians, even though gynecologists are usually the main healthcare providers during the pregnancy (68% according to Descamps et al.)[30]. It is most likely due to the fact that family physicians are traditionally in charge of immunizations.

4.2 Comparison with data from the literature

In Europe, the European Centre for Disease prevention and Control (ECDC) regularly compares the coverage rate in member states. In 2016-2017 the IVC was known for 9 member states, and ranged from 0.5% in Slovenia to 58.6% in the UK (median 25.0%) [39]. In other northern countries similar rates were found, 48.1% in the USA [40], and 30.8% in New Zealand in 2018 [38].

Most data available in France are based on surveys and range from 7% to 26%.

In 2014, Gaudelus et al. found an IVC in pregnant women of 7% through an online survey based on self-administered questionnaires [27]. In 2015, Loubet et al. found a IVC of 26% through an online survey, with a small non-representative sample and many selection bias [29]. Then Descamps et al. calculated an IVC of 7.4% based on the National Perinatal Survey, a large national study conducted in March 2016 [30]. This data was obtained through a declarative method where women were interviewed by a healthcare professional in the postpartum ward before discharge. Our study finds a IVC of 2.4% for women who gave birth in the same period of time (vs 7.4%) and part of this difference can be explained by a social desirability bias [41] and recall bias that may over-estimate the IVC in any declarative vaccination evaluation. Hu et al compared survey data and administrative data in children's vaccination, to evaluate the impact of recall bias by their mother. The results showed the frequency of over-reporting of vaccination in the survey data, up to 6.6% for DPT vaccine, linked to social desirability bias [42].

4.3 Influenza vaccination determinants

The determinants of low IVC in pregnant women have been well studied and two main obstacles to the vaccination are identified. The first obstacle is the lack of proposition from healthcare providers worldwide. In France, as in many other countries, only 25% of women are offered the vaccination [27,43–46], when Canada or the United States offers it to 75% of women [47,48]. It is often linked to the vaccination status of the healthcare provider, that is also known to be very low [49,50]. The second obstacle is the refusal of the vaccine by pregnant women who either fear an impact of the vaccination on the pregnancy or underestimate the gravity of severe influenza. The refusal rate in France is 68% [27,30].

The education of healthcare professional is key to improve IVC in pregnant women [44]. Psarris et al. led a study in Greece in 2018 where the proposition rate rose from 27% to 100% and the IVC from 14% to 94% after a simple information campaign towards healthcare professionals [43].

Another determinant is the availability of the vaccine. Alessandrini et al. led a cross-sectional multicenter study in three maternities in Paris, where vaccination was available immediately during prenatal consultations, free of charge. The coverage increased from 0% to 35.4% [51].

4.4 Strengths and limitations of the study

The main strengths of the study are the size of the study population and the reproducibility of the calculation of the IVC through the EGB database. Our administrative method is reliable to assess a vaccination coverage in a limited time, on a large representative sample, with less bias than a declarative study. Our IVC results on patients over 65 years old are similar to the French official data, validating the use of the database to evaluate the IVC.

Its limitations are those of the database, that can only record reimbursed vaccines, leaving out over the counter purchase and workplace vaccination. Influenza vaccination is recommended for patients over 65 years old, with chronic disease, obese patients, and pregnant women and their vaccine is supposed to be prescribed and fully reimbursed. In addition, anyone can buy it over the counter, for a reasonable price. We assumed that a pregnant woman would turn to her physician before making any health decision during her pregnancy and would have the prescription for the vaccine. However, women who purchased the vaccine over the counter on their own initiative or got the injection in their workplace would have been falsely classified as "unvaccinated" in our study.

Even though our data cannot be fully exhaustive due to this easy access to the vaccine outside of the traditional health care pathway, our results are useful for analysis and comparison purposes.

We identified some limitations in our secondary endpoints:

Pre-existing medical condition were assessed through the existence of a long-term disease data claim, or hospital discharge quotation of an act or a pathology, as well a medication prescribed related to the pathology. We may have overlooked woman that had a condition that put them at risk of a severe influenza, but who requires no specific treatment and had no record of it in their files.

Parity was estimated with the data available in EGB and is indicative. The data collected goes back to 2005, and any pregnancy occurring before that date will not have been taken into account, as well as any prior pregnancy that may have taken place abroad.

We chose to define socioeconomic deprivation by the CMUc affiliation, a specific health insurance with full coverage for low-income patients. However, socioeconomic deprivation is

a wider concept that includes criteria such as insecure employment, lack of familial support, and reduced access to culture, sport or vacation [52]. The EPICES score (Evaluation of precariousness and Inequalities in Health Examination Centers) is more reliable to describe socioeconomic deprivation [53,54]. Unfortunately, it requires data that are not available on the EGB database.

4.5 Perspectives for research and healthcare

The IVC could be improved by information campaigns towards all healthcare professionals involved with pregnancy care, and towards women in general. A national information campaign was issued on French television to promote influenza vaccination during the winter of 2020-2021, and pregnant women were listed as a priority. The impact of such campaigns could be monitored, by issuing IVC data yearly.

Our evaluation of IVC in pregnant women with the EGB database may lead the way to other vaccination evaluations, such as pertussis vaccination coverage. A pertussis booster is often needed to properly protect the newborn from a severe infection, and an evaluation of the vaccination coverage could help improve this protection.

Finally, from the beginning of the COVID-19 pandemic, pregnant women were considered at high-risk of severe forms of the disease, and they were included in specific clinical trial to demonstrate the mRNA vaccine safety and efficacy [55]. The vaccination, while limited at first to women with comorbidities, or with high-exposure professions, was then recommended for all pregnant women in their 2nd and 3rd trimester. The large-scale communication towards healthcare professionals on the necessity of COVID-19 vaccination during pregnancy may have a positive impact on the IVC in the future.

5 Conclusion

The influenza vaccination coverage in pregnant women has significantly increased in France since the 2012 WHO recommendation, particularly during the 2nd and 3rd trimester of pregnancy, but remains very low (4.1% in 2018). Older women and women with a pre-existing condition at risk for severe influenza were more likely to be vaccinated whereas socioeconomic deprived women had a lower coverage. Parity and multifetal pregnancy did not influence the vaccination behavior. The influenza vaccine was mainly prescribed by family physicians. The influenza vaccination needs to be generalized during pregnancy, and a communication campaign towards women and healthcare professionals would be a good start.

Conclusion

La couverture vaccinale antigrippale chez la femme enceinte a augmenté significativement depuis la recommandation de l'OMS en 2012, particulièrement aux deuxièmes et troisièmes trimestres de grossesse, mais reste trop bas (4,1% en 2018). Les femmes plus âgées, et présentant des comorbidités à risque de forme grave de grippe avaient une meilleure couverture vaccinale, alors que les femmes précaires étaient en moyenne moins vaccinées. La parité et les grossesses multiples n'avait pas d'impact sur la couverture vaccinale. Le vaccin antigrippal était prescrit en majorité par les médecins généralistes. La vaccination antigrippale devrait être généralisée pendant la grossesse, et pour cela, une campagne d'information à destination des femmes enceintes et des professionnels de santé pourrait être proposée.

Clermont-Ferrand, le

Pierre CLAVELOU Doyen - Directeur Vu, le Président du Jury de Thèse, Pr Olivier LESENS

2 06.2021

(Date et Signature)

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7 Supplementary material

7.1 Table 1 – International Classification of Diseases (ICD-10) Codes for delivery

060	Preterm labor and delivery
0601	Preterm labor with spontaneous preterm delivery
0603	Preterm delivery without spontaneous labor
067	Labor and delivery complicated by intrapartum hemorrhage, not elsewhere
	classified
068	Labor and delivery complicated by fetal distress
0680	Labor and delivery complicated by fetal heart rate anomaly
0681	Labor and delivery complicated by meconium in amniotic fluid
0682	Labor and delivery complicated by fetal heart rate anomaly and meconium in
	amniotic fluid
0683	Labor and delivery complicated by biochemical evidence of fetal stress
O688	Labor and delivery complicated by other evidence of fetal stress
O689	Labor and delivery complicated by fetal stress, unspecified
069	Labor and delivery complicated by umbilical cord complications
0690	Labor and delivery complicated by prolapse of cord
0691	Labor and delivery complicated by cord around neck, with compression
0692	Labor and delivery complicated by other cord entanglement, with compression
0693	Labor and delivery complicated by short cord
0694	Labor and delivery complicated by vasa previa
0695	Labor and delivery complicated by vascular lesion of cord
0698	Labor and delivery complicated by other cord complications
0699	Labor and delivery complicated by cord complication, unspecified
0755	Delayed delivery after artificial rupture of membranes
0756	Delayed delivery after spontaneous or unspecified rupture of membranes
0757	Vaginal delivery following previous cesarean section
080	Encounter for full-term uncomplicated delivery
0800	Single vertex delivery
0801	Single breech delivery
O808	Other single spontaneous delivery
O809	Single spontaneous delivery, unspecified
081	Single delivery by forceps and vacuum extractor
0810	Low forceps delivery
0811	Mid cavity forceps delivery
O812	Mid cavity forceps V with rotation
0813	Other and unspecified forceps delivery
O814	Vacuum extractor delivery
0815	Delivery by combination of forceps and vacuum extractor
082	Single delivery by cesarean section
O820	Delivery by elective cesarean section
0821	Delivery by emergency cesarean section
0822	Delivery by cesarean section with hysterectomy

0000	
0828	Other single delivery by cesarean section
0829	Delivery by cesarean delivery, unspecified
O83	Other assisted single delivery
O830	Breech extraction
0831	Other assisted breech extraction
0832	Other manipulation-assisted delivery
O833	Delivery of viable fetus in abdominal pregnancy
0834	Destructive operation for delivery
O838	Other specified assisted single delivery
0839	Assisted single delivery, unspecified
O84	Multiple delivery
O840	Multiple delivery, all spontaneous
O841	Multiple delivery, all by forceps and vacuum extractor
O842	Multiple delivery, all by cesarean section
O848	Other multiple delivery
O849	Multiple delivery, unspecified
Z370	Single live birth
Z371	Single stillbirth
Z372	Twins, both liveborn
Z373	Twins, one liveborn, one stillborn
Z374	Twins, both stillborn
Z375	Other multiple births, all liveborn
Z376	Other multiple births, some liveborn
Z377	Other multiple births, all stillborn
Z379	Outcome of delivery, unspecified

7.2 Table 2 - CCAM acts (French classification for clinical procedures) for delivery

JQGA002 Delivery by elective cesarean section, by laparotomy JQGA003 Delivery by cesarean section during labor by laparotomy JQGA004 Delivery by emergency cesarean section by laparotomy JQGA005 Delivery by vaginal cesarean section JQGD001 Single breech delivery JQGD002 Multiple delivery, primiparous JQGD003 Single breech delivery with minimal extraction, primiparous JQGD004 Single breech delivery, primiparous JQGD005 Single breech delivery, multiparous JQGD007 Multiple delivery, multiparous JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous		
JQGA004 Delivery by emergency cesarean section by laparotomy JQGA005 Delivery by vaginal cesarean section JQGD001 Single breech delivery JQGD002 Multiple delivery, primiparous JQGD003 Single breech delivery with minimal extraction, primiparous JQGD004 Single breech delivery, primiparous JQGD005 Single breech delivery, multiparous JQGD007 Multiple delivery, multiparous JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGA002	Delivery by elective cesarean section, by laparotomy
JQGA005 Delivery by vaginal cesarean section JQGD001 Single breech delivery JQGD002 Multiple delivery, primiparous JQGD003 Single breech delivery with minimal extraction, primiparous JQGD004 Single breech delivery, primiparous JQGD005 Single breech delivery, multiparous JQGD007 Multiple delivery, multiparous JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGA003	Delivery by cesarean section during labor by laparotomy
JQGD001 Single breech delivery JQGD002 Multiple delivery, primiparous JQGD003 Single breech delivery with minimal extraction, primiparous JQGD004 Single breech delivery, primiparous JQGD005 Single breech delivery, multiparous JQGD007 Multiple delivery, multiparous JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGA004	Delivery by emergency cesarean section by laparotomy
JQGD002 Multiple delivery, primiparous JQGD003 Single breech delivery with minimal extraction, primiparous JQGD004 Single breech delivery, primiparous JQGD005 Single breech delivery, multiparous JQGD007 Multiple delivery, multiparous JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGA005	Delivery by vaginal cesarean section
JQGD003 Single breech delivery with minimal extraction, primiparous JQGD004 Single breech delivery, primiparous JQGD005 Single breech delivery, multiparous JQGD007 Multiple delivery, multiparous JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGD001	Single breech delivery
JQGD004 Single breech delivery, primiparous JQGD005 Single breech delivery, multiparous JQGD007 Multiple delivery, multiparous JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGD002	Multiple delivery, primiparous
JQGD005 Single breech delivery, multiparous JQGD007 Multiple delivery, multiparous JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGD003	Single breech delivery with minimal extraction, primiparous
JQGD007 Multiple delivery, multiparous JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGD004	Single breech delivery, primiparous
JQGD008 Single breech delivery with minimal extraction, multiparous JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGD005	Single breech delivery, multiparous
JQGD010 Single vertex delivery, primiparous JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGD007	Multiple delivery, multiparous
JQGD011 Manipulation-assisted breech delivery JQGD012 Single vertex delivery, multiparous	JQGD008	Single breech delivery with minimal extraction, multiparous
JQGD012 Single vertex delivery, multiparous	JQGD010	Single vertex delivery, primiparous
	JQGD011	Manipulation-assisted breech delivery
	JQGD012	Single vertex delivery, multiparous
JQGD013 Single breech delivery with maximal extraction, primiparous	JQGD013	Single breech delivery with maximal extraction, primiparous

7.3 Table 3 - ICD codes for non-inclusion criterias

Z640	Unwanted pregnancy
Z33	Denial of pregnancy

7.4 Table 4 - ICD codes for secondary endpoints

Z641	Problems related to multiparity
Z354	Supervision of pregnancy with grand multiparity

7.5 Table 5 - ICD-10 Codes CIM for medical risk factors

7.5.1 Chronic pulmonary diseases

J45	Asthma
J450	Predominantly allergic asthma
J451	Nonallergic asthma
J458	Mixed asthma
J459	Other and unspecified asthma
Z870	Personal history of diseases of the respiratory system
0995	Diseases of the respiratory system complicating pregnancy, childbirth, and the
	puerperium
J961	Chronic respiratory failure
J441	Chronic obstructive pulmonary disease with acute exacerbation, unspecified
J448	Other specified chronic obstructive pulmonary disease
J969	Respiratory failure, unspecified
P279	Unspecified chronic respiratory disease originating in the perinatal period
J40	Bronchitis, not specified as acute or chronic
J410	Simple chronic bronchitis
J411	Mucopurulent chronic bronchitis
J418	Mixed simple and mucopurulent chronic bronchitis
J42	Unspecified chronic bronchitis
J448	Other specified chronic obstructive pulmonary disease
J670	Hypersensitivity pneumonitis due to organic dust
J684	Chronic respiratory conditions due to chemicals, gases, fumes, and vapors
M954	Acquired deformity of chest and rib
Q768	Other congenital malformations of bony thorax
Q769	Congenital malformation of bony thorax, unspecified
Q32	Congenital malformations of trachea and bronchus
Q33	Congenital malformation of lung

Q330	Congenital cystic lung
J47	Bronchiectasis
J980	Diseases of bronchus, not elsewhere classified
P271	Bronchopulmonary dysplasia originating in the perinatal period
Q336	Hypoplasia and dysplasia of lung
E84	Cystic fibrosis
E840	Cystic fibrosis with pulmonary manifestations
E848	Cystic fibrosis with other manifestations
E849	Cystic fibrosis, unspecified

7.5.2 Cardiac diseases

Q20 Congenital malformations of cardiac chambers and connections Q26 Congenital malformations of great veins Q231 Congenital insufficiency of aortic valve Q248 Other specified congenital malformations of heart Q249 Congenital malformation of heart, unspecified I110 Hypertensive heart disease with (congestive) heart failure I130 Hypertensive heart and renal disease with (congestive) heart failure I1300 Hypertensive heart and renal disease with heart failure, LVEF ≥ 50 I1302 Hypertensive heart and renal disease with both (congestive) heart failure and renal failure I1309 Hypertensive heart and renal disease, unspecified I1320 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 I1321 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and < 50		
Q26 Congenital malformations of great veins Q231 Congenital insufficiency of aortic valve Q248 Other specified congenital malformations of heart Q249 Congenital malformation of heart, unspecified I110 Hypertensive heart disease with (congestive) heart failure I130 Hypertensive heart and renal disease with (congestive) heart failure I1300 Hypertensive heart and renal disease with heart failure, LVEF ≥ 50 I1302 Hypertensive heart and renal disease with both (congestive) heart failure and renal failure I1309 Hypertensive heart and renal disease, unspecified I1320 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 I1321 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and < 50 I1322 Hypertensive heart and renal disease with heart and renal failure, LVEF < 40 I1329 Hypertensive heart and renal disease with heart and renal failure, LVEF unspecified I25 Chronic ischemic heart disease I251 Atherosclerotic heart disease I252 Old myocardial infarction I253 Aneurysm of heart I254 Coronary artery aneurysm and dissection I255 Ischemic cardiomyopathy I256 Silent myocardial ischemia I258 Other forms of chronic ischemic heart disease I259 Chronic ischemic heart disease, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	Q20	Congenital malformations of cardiac chambers and connections
Q231 Congenital insufficiency of aortic valve Q248 Other specified congenital malformations of heart Q249 Congenital malformation of heart, unspecified I110 Hypertensive heart disease with (congestive) heart failure I130 Hypertensive heart and renal disease with (congestive) heart failure I1300 Hypertensive heart and renal disease with both (congestive) heart failure and renal failure I1301 Hypertensive heart and renal disease, unspecified I1320 Hypertensive heart and renal disease, unspecified I1321 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 I1321 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and <50	Q209	Congenital malformation of cardiac chambers and connections, unspecified
Other specified congenital malformations of heart Q249 Congenital malformation of heart, unspecified I110 Hypertensive heart disease with (congestive) heart failure I130 Hypertensive heart and renal disease with (congestive) heart failure I130 Hypertensive heart and renal disease with heart failure, LVEF ≥ 50 I1302 Hypertensive heart and renal disease with both (congestive) heart failure and renal failure I1309 Hypertensive heart and renal disease, unspecified I1320 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 I1321 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and <50 I1322 Hypertensive heart and renal disease with heart and renal failure, LVEF < 40 I1329 Hypertensive heart and renal disease with heart and renal failure, LVEF unspecified I25 Chronic ischemic heart disease I251 Atherosclerotic heart disease I252 Old myocardial infarction I253 Aneurysm of heart I254 Coronary artery aneurysm and dissection I255 Ischemic cardiomyopathy I256 Silent myocardial ischemia I258 Other forms of chronic ischemic heart disease I259 Chronic ischemic heart disease, unspecified I499 Cardiac arrhythmia, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	Q26	Congenital malformations of great veins
Q249 Congenital malformation of heart, unspecified I110 Hypertensive heart disease with (congestive) heart failure I130 Hypertensive heart and renal disease with (congestive) heart failure I1300 Hypertensive heart and renal disease with heart failure, LVEF ≥ 50 I1301 Hypertensive heart and renal disease with both (congestive) heart failure and renal failure I1309 Hypertensive heart and renal disease, unspecified I1320 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 I1321 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and <50 I1322 Hypertensive heart and renal disease with heart and renal failure, LVEF <40 I1329 Hypertensive heart and renal disease with heart and renal failure, LVEF unspecified I25 Chronic ischemic heart disease I251 Atherosclerotic heart disease I252 Old myocardial infarction I253 Aneurysm of heart I254 Coronary artery aneurysm and dissection I255 Ischemic cardiomyopathy I256 Silent myocardial ischemia I258 Other forms of chronic ischemic heart disease I259 Chronic ischemic heart disease, unspecified I499 Cardiac arrhythmia, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	Q231	Congenital insufficiency of aortic valve
1110 Hypertensive heart disease with (congestive) heart failure 1130 Hypertensive heart and renal disease with (congestive) heart failure 11300 Hypertensive heart and renal disease with heart failure, LVEF ≥ 50 11302 Hypertensive heart and renal disease with both (congestive) heart failure and renal failure 11309 Hypertensive heart and renal disease, unspecified 11320 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 11321 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and <50	Q248	Other specified congenital malformations of heart
1130 Hypertensive heart and renal disease with (congestive) heart failure 11300 Hypertensive heart and renal disease with heart failure, LVEF ≥ 50 11302 Hypertensive heart and renal disease with both (congestive) heart failure and renal failure 11309 Hypertensive heart and renal disease, unspecified 11320 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 11321 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and <50	Q249	Congenital malformation of heart, unspecified
I1300 Hypertensive heart and renal disease with heart failure, LVEF ≥ 50 I1302 Hypertensive heart and renal disease with both (congestive) heart failure and renal failure I1309 Hypertensive heart and renal disease, unspecified I1320 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 I1321 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and <50	1110	Hypertensive heart disease with (congestive) heart failure
Hypertensive heart and renal disease with both (congestive) heart failure and renal failure Hypertensive heart and renal disease, unspecified Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and <50 Hypertensive heart and renal disease with heart and renal failure, LVEF <40 Hypertensive heart and renal disease with heart and renal failure, LVEF unspecified Chronic ischemic heart disease Chronic ischemic heart disease Did myocardial infarction Aneurysm of heart Coronary artery aneurysm and dissection Silent myocardial ischemia Chronic ischemic heart disease Chronic ischemic heart disease Chronic ischemic heart disease, unspecified Alcoholic cardiomyopathy Cardiac arrhythmia, unspecified Alcoholic cardiomyopathy due to drugs and other external agents	I130	Hypertensive heart and renal disease with (congestive) heart failure
renal failure 11309	I1300	Hypertensive heart and renal disease with heart failure, LVEF ≥ 50
 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50 Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and <50 Hypertensive heart and renal disease with heart and renal failure, LVEF <40 Hypertensive heart and renal disease with heart and renal failure, LVEF unspecified Chronic ischemic heart disease Atherosclerotic heart disease Old myocardial infarction Aneurysm of heart Coronary artery aneurysm and dissection Ischemic cardiomyopathy Silent myocardial ischemia Other forms of chronic ischemic heart disease Chronic ischemic heart disease, unspecified Alcoholic cardiomyopathy Cardiac arrhythmia, unspecified Alcoholic cardiomyopathy due to drugs and other external agents 	l1302	, , ,
Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40, and <50 Hypertensive heart and renal disease with heart and renal failure, LVEF <40 Hypertensive heart and renal disease with heart and renal failure, LVEF unspecified Chronic ischemic heart disease Chronic ischemic heart disease Dischemic heart disease Cold myocardial infarction Aneurysm of heart Coronary artery aneurysm and dissection Isshemic cardiomyopathy Silent myocardial ischemia Other forms of chronic ischemic heart disease Chronic ischemic heart disease, unspecified Alcoholic cardiomyopathy Idea Alcoholic cardiomyopathy Idea Cardiomyopathy due to drugs and other external agents	11309	Hypertensive heart and renal disease, unspecified
I1322 Hypertensive heart and renal disease with heart and renal failure, LVEF <40 I1329 Hypertensive heart and renal disease with heart and renal failure, LVEF unspecified I25 Chronic ischemic heart disease I251 Atherosclerotic heart disease I252 Old myocardial infarction I253 Aneurysm of heart I254 Coronary artery aneurysm and dissection I255 Ischemic cardiomyopathy I256 Silent myocardial ischemia I258 Other forms of chronic ischemic heart disease I259 Chronic ischemic heart disease, unspecified I499 Cardiac arrhythmia, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	I1320	Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 50
Hypertensive heart and renal disease with heart and renal failure, LVEF <40 Hypertensive heart and renal disease with heart and renal failure, LVEF unspecified Chronic ischemic heart disease Chronic ischemic heart disease Chronic ischemic heart disease Cold myocardial infarction Aneurysm of heart Coronary artery aneurysm and dissection Ischemic cardiomyopathy Silent myocardial ischemia Other forms of chronic ischemic heart disease Cardiac arrhythmia, unspecified Alcoholic cardiomyopathy Cardiomyopathy due to drugs and other external agents	I1321	Hypertensive heart and renal disease with heart and renal failure, LVEF ≥ 40,
Hypertensive heart and renal disease with heart and renal failure, LVEF unspecified 125 Chronic ischemic heart disease 1251 Atherosclerotic heart disease 1252 Old myocardial infarction 1253 Aneurysm of heart 1254 Coronary artery aneurysm and dissection 1255 Ischemic cardiomyopathy 1256 Silent myocardial ischemia 1258 Other forms of chronic ischemic heart disease 1259 Chronic ischemic heart disease, unspecified 1499 Cardiac arrhythmia, unspecified 1426 Alcoholic cardiomyopathy 1427 Cardiomyopathy due to drugs and other external agents		and <50
unspecified 125 Chronic ischemic heart disease 1251 Atherosclerotic heart disease 1252 Old myocardial infarction 1253 Aneurysm of heart 1254 Coronary artery aneurysm and dissection 1255 Ischemic cardiomyopathy 1256 Silent myocardial ischemia 1258 Other forms of chronic ischemic heart disease 1259 Chronic ischemic heart disease, unspecified 1499 Cardiac arrhythmia, unspecified 1426 Alcoholic cardiomyopathy 1427 Cardiomyopathy due to drugs and other external agents	I1322	Hypertensive heart and renal disease with heart and renal failure, LVEF <40
I25 Chronic ischemic heart disease I251 Atherosclerotic heart disease I252 Old myocardial infarction I253 Aneurysm of heart I254 Coronary artery aneurysm and dissection I255 Ischemic cardiomyopathy I256 Silent myocardial ischemia I258 Other forms of chronic ischemic heart disease I259 Chronic ischemic heart disease, unspecified I499 Cardiac arrhythmia, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	11329	Hypertensive heart and renal disease with heart and renal failure, LVEF
1251 Atherosclerotic heart disease 1252 Old myocardial infarction 1253 Aneurysm of heart 1254 Coronary artery aneurysm and dissection 1255 Ischemic cardiomyopathy 1256 Silent myocardial ischemia 1258 Other forms of chronic ischemic heart disease 1259 Chronic ischemic heart disease, unspecified 1499 Cardiac arrhythmia, unspecified 1426 Alcoholic cardiomyopathy 1427 Cardiomyopathy due to drugs and other external agents		unspecified
1252 Old myocardial infarction 1253 Aneurysm of heart 1254 Coronary artery aneurysm and dissection 1255 Ischemic cardiomyopathy 1256 Silent myocardial ischemia 1258 Other forms of chronic ischemic heart disease 1259 Chronic ischemic heart disease, unspecified 1499 Cardiac arrhythmia, unspecified 1426 Alcoholic cardiomyopathy 1427 Cardiomyopathy due to drugs and other external agents	125	Chronic ischemic heart disease
I253 Aneurysm of heart I254 Coronary artery aneurysm and dissection I255 Ischemic cardiomyopathy I256 Silent myocardial ischemia I258 Other forms of chronic ischemic heart disease I259 Chronic ischemic heart disease, unspecified I499 Cardiac arrhythmia, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	I251	Atherosclerotic heart disease
1254 Coronary artery aneurysm and dissection 1255 Ischemic cardiomyopathy 1256 Silent myocardial ischemia 1258 Other forms of chronic ischemic heart disease 1259 Chronic ischemic heart disease, unspecified 1499 Cardiac arrhythmia, unspecified 1426 Alcoholic cardiomyopathy 1427 Cardiomyopathy due to drugs and other external agents	1252	Old myocardial infarction
I255 Ischemic cardiomyopathy I256 Silent myocardial ischemia I258 Other forms of chronic ischemic heart disease I259 Chronic ischemic heart disease, unspecified I499 Cardiac arrhythmia, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	1253	Aneurysm of heart
I256 Silent myocardial ischemia I258 Other forms of chronic ischemic heart disease I259 Chronic ischemic heart disease, unspecified I499 Cardiac arrhythmia, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	1254	Coronary artery aneurysm and dissection
Other forms of chronic ischemic heart disease I259 Chronic ischemic heart disease, unspecified I499 Cardiac arrhythmia, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	1255	Ischemic cardiomyopathy
1259 Chronic ischemic heart disease, unspecified 1499 Cardiac arrhythmia, unspecified 1426 Alcoholic cardiomyopathy 1427 Cardiomyopathy due to drugs and other external agents	1256	Silent myocardial ischemia
I499 Cardiac arrhythmia, unspecified I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	1258	Other forms of chronic ischemic heart disease
I426 Alcoholic cardiomyopathy I427 Cardiomyopathy due to drugs and other external agents	1259	Chronic ischemic heart disease, unspecified
I427 Cardiomyopathy due to drugs and other external agents	1499	Cardiac arrhythmia, unspecified
	1426	Alcoholic cardiomyopathy
I50 Heart failure	1427	Cardiomyopathy due to drugs and other external agents
1	150	Heart failure

1500	Congestive heart failure
15000	Congestive heart failure, LVEF ≥ 50
15001	Congestive heart failure, LVEF ≥ 40, and <50
15001	Congestive heart failure, LVEF <40
15002	Congestive heart failure, LVEF unspecified
1501	Left ventricular failure
15010	Left ventricular failure, LVEF ≥ 50
15010	Left ventricular failure, LVEF ≥ 40, and <50
15011	Left ventricular failure, LVEF <40
15012	Left ventricular failure, LVEF v40
1509	Heart failure, unspecified
1517	Cardiomegaly
138	Valvular insufficiency of unspecified valve
1059	Mitral valve disease, unspecified
1069	Rheumatic aortic valve disease, unspecified
1079	Tricuspid valve disease, unspecified
1080	Multiple valve diseases
1081	Disorders of both mitral and tricuspid valves
1082	Disorders of both aortic and tricuspid valves
1083	Combined disorders of mitral, aortic and tricuspid valves
1088	Other multiple valve diseases
1089	Multiple valve disease, unspecified
1091	Rheumatic diseases of endocardium, valve unspecified
1099	Rheumatic heart disease, unspecified
1238	Other current complications following acute myocardial infarction
1348	Other nonrheumatic mitral valve disorders
1349	Nonrheumatic mitral valve disorder, unspecified
1350	Aortic valve stenosis
1359	Aortic valve disorder, unspecified
1379	Pulmonary valve disorder, unspecified
144	Atrioventricular and left bundle-branch block
145	Other conduction disorders
1459	Conduction disorder, unspecified
I471	Supraventricular tachycardia
1472	Ventricular tachycardia
1479	Paroxysmal tachycardia, unspecified
149	Other cardiac arrhythmias
1498	Other specified cardiac arrhythmias
1499	Cardiac arrhythmia, unspecified
1978	Other postprocedural disorders of circulatory system, not elsewhere classified
Q229	Congenital malformation of tricuspid valve, unspecified
Q231	Congenital insufficiency of aortic valve
Q233	Congenital mitral insufficiency
E854	Cardiac amyloidosis
O994	Diseases of the circulatory system complicating pregnancy, childbirth, and the
	puerperium
	Leaster a

O998	Other specified diseases and conditions complicating pregnancy, childbirth, and the puerperium
Z867	Personal history of diseases of the circulatory system
Z95	Presence of cardiac and vascular implants and grafts

7.5.3 Neurological disease

164	Stroke, not specified as hemorrhage or infarction
Z8660	Personal history of diseases of the nervous system and sense organs
G459	Transient cerebral ischemic attack, unspecified
1694	Sequelae of stroke, not specified as hemorrhage or infarction
M63	Disorders of muscle in diseases classified elsewhere
G60	Hereditary and idiopathic neuropathy
G71	Primary disorders of muscles
G710	Muscular dystrophy
G712	Congenital myopathies
G713	Mitochondrial myopathy, not elsewhere classified
G72	Other myopathies
G722	Myopathy due to other toxic agents
G724	Inflammatory myopathy, not elsewhere classified
G728	Other specified myopathies
G729	Myopathy, unspecified
A80	Acute poliomyelitis
A800	Acute paralytic poliomyelitis, vaccine-associated
A801	Acute paralytic poliomyelitis, wild virus, imported
A802	Acute paralytic poliomyelitis, wild virus, indigenous
A803	Acute paralytic poliomyelitis, other and unspecified
A804	Acute nonparalytic poliomyelitis
P358	Congenital poliomyelitis
B91	Sequelae of poliomyelitis
G700	Myasthenia gravis
G702	Congenital and developmental myasthenia
G711	Myotonic disorders
G122	Motor neuron disease
J986	Paralysis of diaphragm
S278	Diaphragm injury

7.5.4 Renal diseases

N009	Acute nephritic syndrome, unspecified
N03	Chronic nephritic syndrome

N031	Chronic nephritic syndrome, focal and segmental glomerular lesions
N032	Chronic nephritic syndrome, diffuse membranous glomerulonephritis
N033	Chronic nephritic syndrome, diffuse mesangial proliferative glomerulonephritis
N034	Chronic nephritic syndrome, diffuse endocapillary proliferative
	glomerulonephritis
N035	Chronic nephritic syndrome, diffuse mesangiocapillary glomerulonephritis
N036	Chronic nephritic syndrome, dense deposit disease
N037	Chronic nephritic syndrome, diffuse crescentic glomerulonephritis
N038	Other glomerulonephritis
N144	Toxic nephropathy, not elsewhere classified
N17	Acute renal failure
N170	Acute renal failure with tubular necrosis
N171	Acute renal failure with acute cortical necrosis
N172	Acute renal failure with medullary necrosis
N18	Chronic kidney disease
N180	End-stage renal disease
N181	Chronic kidney disease, stage 1
N182	Chronic kidney disease, stage 2
N183	Chronic kidney disease, stage 3
N184	Chronic kidney disease, stage 4
N185	Chronic kidney disease, stage 5
N188	Other chronic kidney disease
N189	Chronic kidney disease, unspecified
N19	Unspecified kidney failure
N28	Other disorders of kidney and ureter, not elsewhere classified
N280	Ischemia and infarction of kidney
N288	Other specified disorders of kidney and ureter
N289	Disorder of kidney and ureter, unspecified
N29	Other disorders of kidney and ureter in diseases classified elsewhere
N298	Other disorders of kidney and ureter in other diseases classified elsewhere
N990	Postprocedural renal failure
I120	Hypertensive renal disease with renal failure
I129	Hypertensive renal disease without renal failure
I139	Hypertensive heart and renal disease, unspecified
K767	Hepatorenal syndrome
N040	Nephrotic syndrome with minor glomerular abnormality
N042	Nephrotic syndrome, diffuse membranous glomerulonephritis
N043	Nephrotic syndrome, diffuse mesangial proliferative glomerulonephritis
N044	Nephrotic syndrome, diffuse endocapillary proliferative glomerulonephritis
N045	Nephrotic syndrome, diffuse mesangiocapillary glomerulonephritis
N046	Nephrotic syndrome, dense deposit disease
N047	Nephrotic syndrome, diffuse crescentic glomerulonephritis
N048	Nephrotic syndrome, other morphological lesions
N049	Nephrotic syndrome, unspecified

7.5.5 Sickle cell disease

D57	Sickle-cell disorders
D570	Sickle-cell anemia with crisis
D571	Sickle-cell anemia without crisis
D572	Double heterozygous sickling disorders
D573	Sickle-cell trait
D578	Other sickle-cell disorders

7.5.6 Diabetes

E10	Type 1 diabetes mellitus
E108	Type 1 diabetes mellitus with unspecified complications
E109	Type 1 diabetes mellitus without complications
E11	Type 2 diabetes mellitus
E118	Type 2 diabetes mellitus with unspecified complications
E119	Type 2 diabetes mellitus without complications
E14	Unspecified diabetes mellitus
E148	Unspecified diabetes mellitus with unspecified complications
E149	Unspecified diabetes mellitus without complications

7.5.7 Immunodeficiency

D802	Selective deficiency of immunoglobulin A [IgA]
D803	Selective deficiency of immunoglobulin G [IgG] subclasses
D808	Other immunodeficiencies with predominantly antibody defects
D81	Combined immunodeficiencies
D811	Severe combined immunodeficiency [SCID] with low T- and B-cell numbers
D812	Severe combined immunodeficiency [SCID] with low or normal B-cell numbers
D813	Adenosine deaminase [ADA] deficiency
D816	Major histocompatibility complex class I deficiency
D817	Major histocompatibility complex class II deficiency
D818	Other combined immunodeficiencies
D819	Combined immunodeficiency, unspecified
D822	Immunodeficiency with short-limbed stature
D828	Immunodeficiency associated with other specified major defects
D830	Common variable immunodeficiency
D831	Common variable immunodeficiency with predominant immunoregulatory T-
	cell disorders

D832	Common variable immunodeficiency with autoantibodies to B- or T-cells
D838	Other common variable immunodeficiencies
D839	Common variable immunodeficiency, unspecified
D848	Other specified immunodeficiencies
D849	Immunodeficiency, unspecified
Z94802	Allogeneic hematopoietic stem-cell transplantation, bone marrow transplant,
	related donor
Z94804	Allogeneic hematopoietic stem-cell transplantation, bone marrow transplant,
	unrelated donor
Z940	Kidney transplant status
Z941	Heart transplant status
Z942	Lung transplant status
Z943	Heart and lungs transplant status
Z944	Liver transplant status
Z945	Skin transplant status
Z946	Bone transplant status
Z947	Corneal transplant status
Z948	Other transplanted organ and tissue status
Z94802	Allogeneic hematopoietic stem-cell transplantation, bone marrow transplant,
	related donor
Z94804	Allogeneic hematopoietic stem-cell transplantation, bone marrow transplant,
	unrelated donor
Z949	Transplanted organ and tissue status, unspecified
B24	Unspecified human immunodeficiency virus [HIV] disease
B240	Unspecified pre-acquired immunodeficiency syndrome [AIDS]
B241	Unspecified full-blown AIDS
B249	Unspecified HIV infection
•	<u> </u>

7.5.8 Hepatic diseases

B180	Chronic viral hepatitis B with delta-agent
B181	Chronic viral hepatitis B without delta-agent
B182	Chronic viral hepatitis C
B189	Chronic viral hepatitis, unspecified
B199	Unspecified viral hepatitis without hepatic coma
C787	Secondary malignant neoplasm of liver and intrahepatic bile duct
D376	Liver, gallbladder, and bile ducts neoplasm
K701	Alcoholic hepatitis
K703	Alcoholic cirrhosis of liver
K704	Alcoholic hepatic failure
K709	Alcoholic liver disease, unspecified
K711	Toxic liver disease with hepatic necrosis
K713	Toxic liver disease with chronic persistent hepatitis

K714 Toxic liver disease with chronic lobular hepatitis K715 Toxic liver disease with chronic active hepatitis K716 Toxic liver disease with hepatitis not elsewhere classified K718 Toxic liver disease with other disorders of liver K719 Toxic liver disease, unspecified K738 Other chronic hepatitis, not elsewhere classified K739 Chronic hepatitis, unspecified K740 Hepatic fibrosis K746 Other and unspecified cirrhosis of liver K754 Autoimmune hepatitis K758 Other specified inflammatory liver diseases K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder Z871 Personal history of diseases of the digestive system		
K716 Toxic liver disease with hepatitis not elsewhere classified K718 Toxic liver disease with other disorders of liver K719 Toxic liver disease, unspecified K738 Other chronic hepatitis, not elsewhere classified K739 Chronic hepatitis, unspecified K740 Hepatic fibrosis K746 Other and unspecified cirrhosis of liver K754 Autoimmune hepatitis K758 Other specified inflammatory liver diseases K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome Other specified perinatal digestive system disorder	K714	Toxic liver disease with chronic lobular hepatitis
K718 Toxic liver disease with other disorders of liver K719 Toxic liver disease, unspecified K738 Other chronic hepatitis, not elsewhere classified K739 Chronic hepatitis, unspecified K740 Hepatic fibrosis K746 Other and unspecified cirrhosis of liver K754 Autoimmune hepatitis K758 Other specified inflammatory liver diseases K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K715	Toxic liver disease with chronic active hepatitis
K719 Toxic liver disease, unspecified K738 Other chronic hepatitis, not elsewhere classified K739 Chronic hepatitis, unspecified K740 Hepatic fibrosis K746 Other and unspecified cirrhosis of liver K754 Autoimmune hepatitis K758 Other specified inflammatory liver diseases K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K716	Toxic liver disease with hepatitis not elsewhere classified
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K739 Chronic hepatitis, unspecified K740 Hepatic fibrosis K746 Other and unspecified cirrhosis of liver K754 Autoimmune hepatitis K758 Other specified inflammatory liver diseases K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K719	Toxic liver disease, unspecified
K740 Hepatic fibrosis K746 Other and unspecified cirrhosis of liver K754 Autoimmune hepatitis K758 Other specified inflammatory liver diseases K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K738	Other chronic hepatitis, not elsewhere classified
K746 Other and unspecified cirrhosis of liver K754 Autoimmune hepatitis K758 Other specified inflammatory liver diseases K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K739	Chronic hepatitis, unspecified
K754 Autoimmune hepatitis K758 Other specified inflammatory liver diseases K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K740	Hepatic fibrosis
K758 Other specified inflammatory liver diseases K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K746	Other and unspecified cirrhosis of liver
K759 Inflammatory liver disease, unspecified K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K754	Autoimmune hepatitis
K768 Other specified diseases of liver K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K758	Other specified inflammatory liver diseases
K769 Liver disease, unspecified K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K759	Inflammatory liver disease, unspecified
K831 Obstruction of bile duct P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K768	Other specified diseases of liver
P591 Inspissated bile syndrome P788 Other specified perinatal digestive system disorder	K769	Liver disease, unspecified
P788 Other specified perinatal digestive system disorder	K831	Obstruction of bile duct
	P591	Inspissated bile syndrome
Z871 Personal history of diseases of the digestive system	P788	Other specified perinatal digestive system disorder
	Z871	Personal history of diseases of the digestive system

7.5.9 Obesity

E6600	Obesity due to excess calories in adults with a body mass index (BMI) ≥30 kg / m2 or more and < 40 kg / m2, or obesity due to excess calories in children
E6601	Obesity due to excess calories in adults with BMI ≥40 kg / m2 and <50 kg / m2
E6602	Obesity due to excess calories in adults with BMI ≥50 kg / m2
E6609	Obesity due to excess calories in adults, BMI not specified
E6610	Drug-induced obesity with BMI ≥30 kg / m2 and < 40 kg / m2, or drug-related
	childhood obesity
E6611	Drug-induced obesity with BMI ≥40 kg / m2 and < 50 kg / m2
E6620	Morbid obesity with alveolar hypoventilation, with BMI ≥30 kg / m2 and < 40 kg / m2
E6690	Unspecified obesity, adult with BMI ≥30 kg / m2 and <40 kg / m2, or unclear childhood obesity
E6691	Unspecified obesity, adult with BMI ≥40 kg / m2 and < 50 kg / m2
E6692	Unspecified adult obesity, with BMI ≥ 50 kg / m2
E6699	Unspecified adult obesity, with BMI not specified

7.6 Table 6 – Long term disease codes for preexisting condition at risk for severe influenza

ALD 1	Stroke
ALD 2	Bone marrow failures and others chronic cytopenias, myelo-dysplastic
	syndromes
ALD 5	Chronic cardiac arrhythmia
ALD 6	Chronic hepatitis and cirrhosis of liver
ALD 7	Severe immunodeficiency, Human Immunodeficiency Virus infection, severe
	immunodeficiency
ALD 8	Type 1 and 2 diabetes
ALD 9	Severe neuromuscular disorders
ALD 10	Sickle cell disease
ALD 13	Ischemic heart disease
ALD 14	Severe respiratory insufficiency due to chronic obstructive pulmonary disease
ALD 18	Cystic fibrosis
ALD 19	Chronic kidney disease
ALD 21	Polyarteritis nodosa, lupus erythematosus, systemic sclerosis
ALD 22	Rheumatoid arthritis
ALD 24	Crohn disease
ALD 25	Multiple sclerosis
ALD 27	Ankylosing spondylitis
ALD 28	Transplanted organ or tissue
ALD 30	Malignant tumor, malignant neoplasm of lymphoid, hematopoietic and related
	tissue

7.7 Table 7 - ATC drugs related to condition at risk for severe influenza

A 10	Drugs used in diabetes
C01	Cardiac therapy
C02	Antihypertensives
C03	Diuretics
C07	Beta blocking agents
C08	Calcium channel blockers
C09	Agents acting on the renin–angiotensin system
L02B	Hormone antagonists and related agents
L04	Immunosuppressants
R03	Drugs for obstructive airway diseases

SERMENT D'HIPPOCRATE (Conseil national de l'ordre des médecins)

Au moment d'être admise à exercer la médecine, je promets et je jure d'être fidèle aux lois de l'honneur et de la probité.

Mon premier souci sera de rétablir, de préserver ou de promouvoir la santé dans tous ses éléments, physiques et mentaux, individuels et sociaux.

Je respecterai toutes les personnes, leur autonomie et leur volonté, sans aucune discrimination selon leur état ou leurs convictions. J'interviendrai pour les protéger si elles sont affaiblies, vulnérables ou menacées dans leur intégrité ou leur dignité. Même sous la contrainte, je ne ferai pas usage de mes connaissances contre les lois de l'humanité.

J'informerai les patients des décisions envisagées, de leurs raisons et de leurs conséquences. Je ne tromperai jamais leur confiance et n'exploiterai pas le pouvoir hérité des circonstances pour forcer les consciences.

Je donnerai mes soins à l'indigent et à quiconque me les demandera. Je ne me laisserai pas influencer par la soif du gain ou la recherche de la gloire.

Admise dans l'intimité des personnes, je tairai les secrets qui me seront confiés. Reçue à l'intérieur des maisons, je respecterai les secrets des foyers et ma conduite ne servira pas à corrompre les mœurs.

Je ferai tout pour soulager les souffrances. Je ne prolongerai pas abusivement les agonies. Je ne provoquerai jamais la mort délibérément.

Je préserverai l'indépendance nécessaire à l'accomplissement de ma mission. Je n'entreprendrai rien qui dépasse mes compétences. Je les entretiendrai et les perfectionnerai pour assurer au mieux les services qui me seront demandés.

J'apporterai mon aide à mes confrères ainsi qu'à leurs familles dans l'adversité.

Que les hommes et mes confrères m'accordent leur estime si je suis fidèle à mes promesses ; que je sois déshonorée et méprisée si j'y manque.

SERMENT D'HIPPOCRATE

En présence des Maîtres de cette FACULTE et de mes chers CONDISCIPLES, je promets et je jure d'être fidèle aux lois de l'Honneur et de la Probité dans l'exercice de la Médecine.

Je donnerai mes soins gratuits à l'indigent et je n'exigerai jamais un salaire au-dessus de mon travail. Admis dans l'intérieur des maisons, mes yeux ne verront pas ce qui s'y passe, ma langue taira les secrets qui me seront confiés et mon état ne servira pas à corrompre les mœurs ni à favoriser le crime.

Respectueux et reconnaissant envers mes MAÎTRES, je rendrai à leurs enfants l'instruction que j'ai reçue de leurs pères.

Que les HOMMES m'accordent leur estime si je suis fidèle à mes promesses. Que je sois couvert d'OPPROBRE et méprisé de mes confrères si j'y manque.

Trends of influenza vaccination coverage in pregnant women, a ten-year analysis from a French healthcare database

Abstract

Background Pregnant women have a high risk of severe influenza, associated with obstetrical complications. The World Health Organization (WHO) has recommended influenza vaccination for all pregnant women since 2012, but the vaccination coverage remains low worldwide, especially in Europe, due to a lack of proposition from the health care providers, and a high refusal rate from the women. The primary aim of this study was to estimate the influenza vaccination coverage (IVC) in a population of pregnant women in France, and to analyse its evolution from 2009 to 2018. The secondary objective was to describe the vaccinated population and to find determinants associated with the vaccination.

Methods This retrospective cohort study is based on the EGB French health care database, a representative sample of the French population containing data from the health insurance system. All pregnant women who delivered medically or spontaneously over the 2009-2018 period were included.

Results In the 2009-2018 period, only 1.2% pregnant women were vaccinated against influenza (n=875/72 207; 95%CI 1.14-1.30). The IVC significantly increased after the 2012 WHO recommendation, from 0.33% to 1.79% (p<0,001) but remained low (4.1% in 2018). Women younger than 25 years old had a low coverage (0.6%) whereas women over 35 years old were more likely to get the influenza vaccine (1.7%; OR: 2.82, 95%CI 2.14-3.71). The vaccination behavior was not influenced by multifetal pregnancy or parity, but socioeconomically deprived women were less likely to be vaccinated (OR: 0.81, 95%CI: 0.67-0.98). Women with pre-existing medical conditions had an overall higher vaccination rate (2.5%; OR: 2.32, 95%CI: 1.94-2.77). The vaccine was mainly prescribed by family physicians (58%).

Conclusion Influenza vaccination in pregnant women in France remains very low, particularly in younger, healthy women, and measures such as information campaigns need to be undertaken to improve the coverage.

Key-words Pregnancy; influenza; vaccination; coverage; EGB

Evolution de la couverture vaccinale antigrippale chez les femmes enceintes de 2009 à 2018 à partir de la base de données EGB

Résumé

CONTEXTE: Les femmes enceintes ont un risque accru de contracter une forme grave d'infection grippale, allant jusqu'à la détresse respiratoire, associé à de nombreuses complications obstétricales. Depuis 2012, l'Organisation Mondiale de la Santé recommande la vaccination antigrippale à toutes les femmes enceintes, quel que soit le terme de leur grossesse, mais la couverture vaccinale reste basse dans le monde dans cette population. Les femmes enceintes refusent souvent le vaccin, mais il est également insuffisamment proposé par les professionnels de santé.

OBJECTIF: L'objectif principal de l'étude était d'évaluer l'évolution de la couverture vaccinale antigrippale chez les femmes enceintes en France sur 10 ans, et de mesurer l'impact de la recommandation de 2012 sur le comportement vaccinal. Les objectifs secondaires étaient de décrire la population vaccinée pour caractériser les éléments en faveur de la vaccination.

METHODE: Cette étude pharmaco-épidémiologique rétrospective a été menée à partir de la base de données EGB (Echantillon Généraliste des Bénéficiaires), et a inclus toutes les femmes ayant accouché sur la période de l'étude, à l'exception des dénis de grossesse et grossesse non désirées (n=72 207). La couverture vaccinale a été calculée à l'aide des données de remboursement des vaccins antigrippaux.

RESULTATS: La couverture vaccinale a augmenté de façon significative de 2009 à 2018 (de 0,33% à 1,79%; p<0,001) mais reste globalement basse (jusqu'à 4,1% en 2018). Les femmes les plus jeunes et celles considérées comme précaires étaient les moins vaccinées, alors que les femmes présentant une comorbidité avaient une couverture vaccinale plus élevée. Le vaccin était majoritairement prescrit par des médecins généralistes (58%).

CONCLUSION: La couverture vaccinale antigrippale reste basse en France malgré les recommandations, particulièrement chez les femmes les plus jeunes, en bonne santé. Des campagnes d'informations auprès des femmes et des professionnels de santé pourraient être entreprises pour améliorer cette couverture vaccinale.

Mots clés: vaccination, grippe, grossesse, EGB, couverture vaccinale.