

Immigration and Adverse Pregnancy Outcomes in an Italian Free Care Hospital

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Introduction: The ever-increasing wave of immigration in Italy has posed demanding challenges in the management of the new multiethnic obstetric population. The aim of this study was to compare pregnancy and perinatal outcomes between immigrants and the native population in an Italian public hospital.

Materials and Methods: Singleton pregnant women (≥ 24 weeks of gestation) who delivered during a 3-year period in an Italian free care hospital were included. Long-term (≥ 2 years of residence) immigrant patients were divided into 4 groups according to their ethnic origin: Europeans, Asians, Latin Americans, and Africans. Perinatal indicators of obstetric outcomes were collected and compared between immigrants and Italians.

Results: Of the 3556 patients included, 1092 were immigrants and 2464 Italians. The immigrant cohort experienced a higher rate of macrosomia (1.8% vs 0.6%; $p = 0.001$), very low birth weight (1.3% vs 0.6%; $p = 0.048$), very early preterm delivery (1.4% vs 0.4%; $p = 0.048$), and gestational diabetes mellitus (1.8% vs 0.5%; $p = 0.003$) compared with the native population. The overall rate of cesarean sections was greater among Italians (56% vs 45.8%; $p < 0.001$). Among ethnic groups, Europeans and Latin Americans reported a higher rate of preterm delivery (20.2% and 19%, respectively; $p < 0.001$). Latin Americans carried also a greater risk of fetal macrosomia (3.6%; $p < 0.008$), while the rate of very low birth weight was higher among Europeans and Africans (2% and 1.8%, respectively; $p < 0.04$).

Conclusion: Obstetricians should pay special attention to the potential disparities in pregnancy outcomes between immigrants and the native population. Future efforts should focus on reducing preterm delivery and glucose dysmetabolism among pregnant immigrants.

Keywords: ethnic disparities, immigration, maternal outcomes, neonatal outcomes, pregnancy

Introduction

The increasing phenomenon of international migration has led to a greater recognition of the ethnic disparities in obstetric and perinatal healthcare. Immigration has been one of the most relevant phenomena characterizing Italian society of the past 40 years. Over the last decades, Italy has faced, as well as other European societies, a huge wave of incoming immigrants, which has led to a progressive change in the baseline socio-demographic and clinical characteristics of obstetric patients. This epidemiologic shift is gradually having obvious implications on the Italian maternal healthcare, posing a substantial clinical challenge for obstetricians.¹

Wide inequities in terms of obstetric complications across different ethnic groups have been well documented in the literature. Immigrant women seem to display a higher prevalence of preeclampsia, gestational diabetes, preterm delivery, low birth

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weight, fetal growth restriction, intrauterine fetal demise, maternal mortality, and cesarean section (CS) rate.^{2–9} The underlying etiopathogenetic mechanisms are poorly understood and likely to be multifactorial, including genetic and epigenetic factors, socio-cultural disparities, poor socio-economic status, inadequate lifestyles and nutrition, and different maternal health behaviors and attitudes.^{10–14} Immigrants often encounter linguistic, socio-economic, and political barriers, resulting eventually in reduced access to the healthcare system and increased perinatal morbidity.^{12–19} However, currently available data are controversial, and it remains unclear whether immigrated mothers have different obstetrical outcomes compared with the native population. Indeed, few studies have described the paradox of better perinatal outcomes among immigrants, the so-called “healthy-migrant effect”, probably linked to a presumed migratory selection of healthy individuals, along with cultural protective factors and social support.²⁰

Italian demographic data showed that the number of native births has significantly decreased over the last 10 years, affecting couples with both native partners more than couples with at least one foreign partner. In 2016, the last Italian demographic report stated that immigrants were approximately 5 million and accounted for 8.6% of the overall Italian population, as opposed to nearly 2 million in 2005. Italian newborns from immigrant women were 77,397.^{20,21}

The ongoing issue of the ethnic disparities in obstetric healthcare is receiving increasing attention and further research is required to address the underlying mechanisms, in order to better understand and manage the increasingly multiethnic population. Recently, several Italian hospitals have focused their research on the obstetric outcomes of immigrated women compared to the native population; however, results were quite heterogeneous.^{22–25}

The aim of the study was to compare pregnancy and perinatal outcomes between long-term (≥ 2 years of residence) immigrant women and the native population in an Italian free care hospital.

Materials and Methods

This is an observational retrospective study including nulliparous or primiparous singleton pregnant women who delivered starting from 24 weeks of gestational age from January 1, 2017 to December 31, 2019 at the Obstetrical Department of Umberto I Hospital of Rome. Our institution is a tertiary level referral center for all obstetric complications with a neonatal intensive care unit (NICU). All patients provided written informed consent

and procedures followed were in accordance with the Helsinki declaration. The present study received ethical approval from the Institutional Review Board of Umberto I Hospital. Demographic and clinical data were retrospectively collected from our hospital admission records.

Only immigrant women who have been resident in Italy for at least 2 years were included. The patients filled in a self-administered multilanguage questionnaire enquiring about sociodemographic data. Spontaneous miscarriages and elective pregnancy terminations were excluded. Twin pregnancies were also excluded to avoid a potential bias in some of the analyzed obstetric indicators, such as birth weight and gestational age at delivery. Immigrant patients were divided into four groups according to their ethnic origin: non-Italian Europeans, Asians, Latin Americans, and Africans. A control group of Italian women admitted to our Department in the same period was provided.

The following perinatal care indicators were evaluated: mode of conception (spontaneous or medically assisted), gestational age (GA) at delivery, birth weight, delivery mode (vaginal delivery versus CS). Obstetric complications were divided into maternal and fetal. Maternal complications included: diabetes mellitus (divided into pregestational, gestational A1 and A2, according to the White classification²⁶), pregnancy-induced hypertension (PIH, including gestational hypertension, preeclampsia, eclampsia, and chronic hypertension), placental abruption, placental adhesive disorders (PAD), and post-partum hemorrhage (PPH). Fetal complications included: (a) pre-term delivery (PTD), defined as a GA at birth <37 weeks, which is subdivided into very early PTD (GA at birth between 24 and 28 weeks), early PTD (GA at birth between 28 and 34 weeks), and late PTD (GA at birth between 34 and 37 weeks); (b) low birth weight (LBW), defined as a birth weight <2500 g regardless of gestational age, and very low birth weight (VLBW) when less than 1500 g; (c) macrosomia, defined as a birth weight ≥ 4500 g.

The incidence rates were analyzed using the Fisher exact test. Normality testing was performed to determine whether data were sampled from a Gaussian distribution. The Chi-square test was used to evaluate the association between two categorical variables. The *t*-test and Mann–Whitney *U*-test were used to compare continuous parametric and nonparametric variables, respectively. Statistical software SPSS version 25.0 was used for all

the analyses. A p value <0.05 was considered statistically significant.

Results

Of the 3556 patients enrolled, 1092 (30.7%) were immigrants (Europeans: 495; Asians: 268; Latin Americans: 110; Africans: 219) and 2464 (69.3%) were Italian. The mean maternal age was 29.31 ± 5.92 years among immigrants and 32.59 ± 5.38 years among native women ($p < 0.05$).

Maternal reproductive and neonatal characteristics are summarized in Table 1. No significant differences were found between the two groups regarding the rate of medically assisted conceptions (8.6% vs 9.5%, respectively) and the GA at delivery (38.74 ± 2.22 weeks vs 38.37 ± 2.50 weeks, respectively). The overall rate of CS was significantly higher in Italians (56% vs 45.8%; $p < 0.001$). The incidence of repeated CS was 30% in Italians and 20% in immigrants. Macrosomia (1.8% vs 0.6%; $p = 0.001$) and VLBW (1.3% vs 0.6%; $p = 0.048$) were significantly higher in immigrants compared to Italian women. Results from the interethnic analysis reported a significantly higher rate of VLBW in Europeans (2.0%) and Africans (1.8%) compared to other groups ($p < 0.04$) and of macrosomia in Latin Americans (3.6%; $p < 0.008$) (Figure 1).

Comparison of pregnancy outcomes between immigrants and native women is summarized in Table 2. Preterm delivery occurred in 16% of immigrants and 13.5% of Italians. Very early PTD was significantly higher in immigrant than Italian women (1.4% vs 0.4%; $p = 0.048$). No differences were found regarding early and late PTD. The subanalysis of PTD among different ethnic groups revealed a significant difference between Europeans and Latin Americans (20.2% vs 19%, respectively) compared with Asians (12.7%), Africans (9.1%), and Italians (13.5%) ($p < 0.01$). No significant differences between immigrants and Italians were reported as regards IUGR (3.1% vs 3.3%, respectively), pregestational diabetes (0.1% vs 0.2%, respectively), pregnancy-induced hypertension (5.1% vs 5.4%, respectively), PAD (1.2% vs 0.9%, respectively), and postpartum hemorrhage (1.5% vs 1.3%, respectively). Finally, gestational diabetes mellitus A2 was significantly higher in immigrants than Italians (1.8% vs 0.5%; $p = 0.03$) (Figure 2). Additional ethnic-group specific data are summarized in Supplemental Table 1.

Discussion

In the present study, we aimed to evaluate the effect of immigration on adverse maternal and perinatal outcomes

Table 1 Maternal, Reproductive and Neonatal Characteristics

Variables	Immigrants (n=1092)	Italians (n=2464)	p value
Maternal age (years)*	29.31 ± 5.92	32.59 ± 5.38	0.048
Mode of conception			NS
Spontaneous	998 (91.4%)	2230 (90.5%)	
ART	94 (8.6%)	234 (9.5%)	
GA at delivery (weeks)*	38.74 ± 2.22	38.37 ± 2.50	NS
Mode of delivery			<0.001
VD	537 (49.2%)	1030 (41.8%)	
OD	55 (5%)	54 (2.2%)	
CS	500 (45.8%)	1380 (56%)	
Birth weight (g)*	3062.19 ± 692.34	3056.94 ± 663.8	NS
< 1500	14 (1.3%)	15 (0.6%)	0.048
1500–2500	175 (16%)	396 (16.1%)	NS
2500–4500	883 (80.9%)	2038 (82.7%)	NS
> 4500	20 (1.8%)	15 (0.6%)	0.001
Apgar score			
1-minute	8.15 ± 1.13	8.12 ± 1.33	NS
5-minute	9.15 ± 1.17	9.27 ± 0.83	NS

Note: *Data are expressed as mean $\pm$ standard deviation (SD).

Abbreviations: ART, assisted reproductive technology; CS, cesarean section; GA, gestational age; NS, not significant; VD, vaginal delivery.

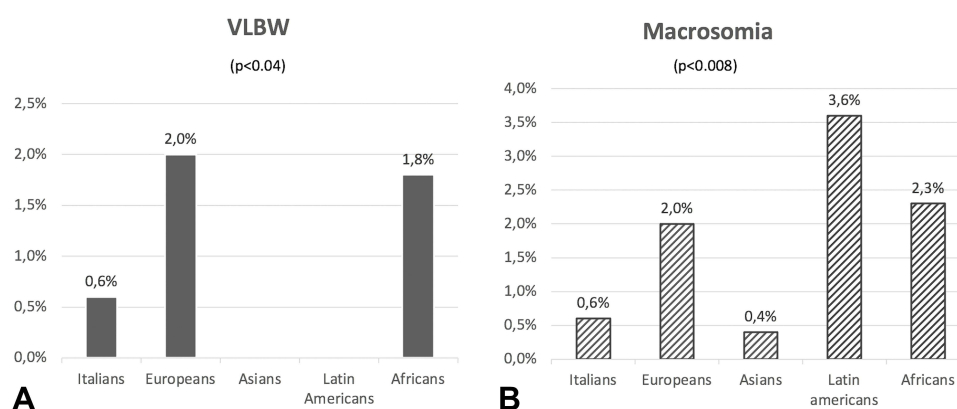


Figure 1 VLBW (A) and macrosomia (B) among different ethnic groups.

in a reference tertiary Italian public hospital. During the study period, immigrant patients requesting obstetric care were 30.1% of all patients admitted as opposed to 10% recorded ten years before. The prevalence was in line with that seen in other European countries, such as Turkish (32%)²² and Germany (32.9%).²³

We analyzed the obstetric outcomes of 1092 immigrants compared to 2464 native patients, with a glimpse into potential differences in pregnancy complications. In Italy, hospitalization, surgical procedures, and necessary medications are provided free of charge for the immigrant population as established by the health system. However,

albeit unlikely, we cannot rule out the financial barrier as a potential underlying risk factor for adverse obstetric outcomes among immigrants. In dealing with this multiethnic obstetric population, we observed that many other factors may have a negative impact on pregnancy outcomes and reduce the quality of care offered, such as poor access to healthcare due to the lack of awareness of its free care nature, delay in looking for prenatal care, missed follow-ups, and communication problems due to the language barrier, among others. An appropriate antenatal care plays a crucial role in facilitating early detection and mitigation of modifiable risk factors during pregnancy.

Table 2 Pregnancy Outcomes in Immigrants and Italians

Variables	Immigrants (n=1092)	Italians (n=2464)	p value
PTD	175 (16%)	332 (13.5%)	NS
Very Early PTD	15 (8.6%)	10 (3%)	0.048
Early PTD	60 (34.3%)	108 (32.5%)	NS
Late PTD	100 (57.1%)	214 (64.5%)	NS
IUGR	34 (3.1%)	81 (3.3%)	NS
Diabetes	39 (3.6%)	63 (2.6%)	NS
GDM A1	18 (46.2%)	44 (69.9%)	NS
GDM A2	20 (51.3%)	13 (20.6%)	0.03
Pregestational	1 (2.6%)	6 (9.5%)	NS
PIH	56 (5.1%)	132 (5.4%)	NS
PAD	13 (1.2%)	23 (0.9%)	NS
Placenta previa	12 (92.3%)	21 (91.3%)	NS
Placenta accreta	1 (7.7%)	2 (8.7%)	NS
Placental abruption	3 (0.3%)	12 (0.5%)	NS
PPH	16 (1.5%)	31 (1.3%)	NS

Abbreviations: GDM A1, gestational diabetes mellitus A1 (White classification class); GDM A2, gestational diabetes mellitus A2 (White classification class); IUGR, intrauterine growth retardation; NS, not significant; PAD, placental adhesive disorders; PIH, pregnancy-induced hypertension; PPH, post-partum hemorrhage; PTD, preterm delivery.

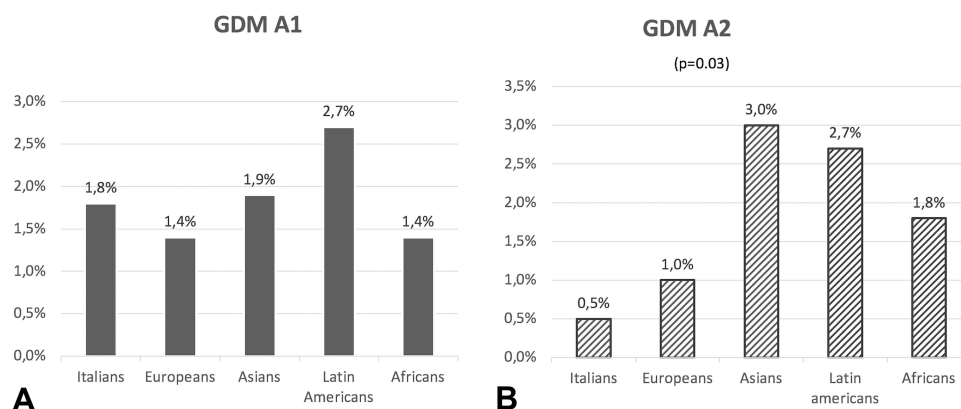


Figure 2 GDM A1 (A) and GDM A2 (B) among different ethnic groups.

Improved communication, comprehension and integration would further support immigrant women in accessing pre-natal health care.

Recently, several Italian studies concerning the obstetric outcomes of immigrated women compared to the native population have been published, albeit with controversial findings.^{24–27} A retrospective Italian study by Rizzo et al found no significant differences between immigrants and native patients with regard to pregnancy outcomes.²⁴ In 2011, Zanconato et al reported significant differences in perinatal outcomes between immigrant and Italian women, the former carrying a higher risk of GA ≤ 32 weeks at delivery, very low birth weight, and cesarean sections.²⁵ Another Italian study by Sosta et al found that preterm delivery occurred more frequently in immigrants, mainly Africans, and was not associated with delayed access to prenatal care.²⁷ Finally, Cacciani et al suggested that perinatal outcomes were worse among infants of immigrant mothers, including very PTD, low Apgar score, and need for NICU.²⁸ Literature on this topic is scant and there is still no consensus on whether there is a difference in terms of obstetric outcomes between native and immigrant mothers. A putative underlying reason of this controversy is that the studies published so far do not consider the duration of the residence in Italy of immigrant women. Indeed, it was interesting to see a comparison between natives and immigrant mothers living in Italy for a minimum of 2 years and still find significant differences between the two groups. However, it should be underlined that many other variables can explain such differences, such as poor access to healthcare, social integration, genetic predisposition, sociocultural and lifestyle factors, and all these should be further addressed in future, multicenter studies.

In our study, the maternal age was higher in Italian patients compared to immigrants and this could be linked to delayed reproductive age in Italy due to socio-economic factors.²⁹ A two-fold increase in the frequency of very early PTD, which is the most severe adverse outcome, was observed in immigrants as compared to Italian women. Increased PTD rates in immigrants have been reported in several Italian studies and the African mothers seem to be the most affected.^{25,27}

The overall incidence of CS in the present study (46–56%) is higher than the one recommended in obstetric care (ie, 5–15% of all births).^{29,30} This could be due to the fact that the rate of vaginal delivery after CS is extremely low in our Department, while repeated CS reach up to 30% of all CS and the rate of primary CS on maternal request is around 8.6%.³¹

Obstetricians should discourage the policy to routinely perform elective CS after a previous one or upon maternal request without an appropriate indication and adequately counsel patients regarding the risks of cesarean sections over vaginal delivery. Controversial data exist in the literature concerning the differences on the CS rate between native and immigrant population. Some studies have found a high rate of CS in the immigrant population.^{32–34} On the other hand, an Italian study by Zanconato et al found that the rate of CS was higher in the native population.²⁵ Further information could derive by an accurate analysis of specific CS indications in both Italian and immigrant group.

Since evaluating obstetric outcomes in a heterogeneous immigrant population has its obvious limitations, we also carried out a subanalysis with each of the 4 immigrant groups. Within our study population, Europeans and Latin Americans experienced a greater risk of PTD. Latin

Americans carried a higher risk of fetal macrosomia, which may be related to poorly controlled gestational diabetes mellitus.³⁵ Finally, the rate of VLBW was significantly higher in Europeans and Africans. In the future, it would be interesting to correlate the degree of integration and the duration of residency in Italy with the risk of developing obstetric complications; good integration and long-term residence may exert a positive impact on health of pregnant women and pregnancy outcomes.

The present study contains both limitations and strengths. Limitations include the following topics: (a) the retrospective study design and the intrinsic risk of confounding; (b) the language barrier, which could have led to an incomplete medical history of immigrant and a difficulty in clinical procedure explanation; (c) the lacking data on the socioeconomic status and education level and quality of prenatal care. Strengths are the high number of the population study and the homogeneity of treatment as all patients were referred to the same medical equipe.

There are still complex gaps in the obstetric management of multiethnic pregnant populations. In our experience, immigration demonstrated to be a major risk factor for several adverse obstetric outcomes, such as very low birth weight, macrosomia, very early preterm delivery, gestational diabetes. Italian health professionals in the field of obstetrics should be more sensitive to the issue of migration medicine as the ethnic background can undoubtedly influence the incidence of maternal-fetal complications. In the era of precision medicine, the obstetric behavior should also consider the ethnic-specific risk of developing pregnancy complications. Future efforts should focus on reducing preterm delivery and glucose dysmetabolism among pregnant immigrants in Italy.

Statement of Ethics

The research was conducted ethically in accordance with the World Medical Association Declaration of Helsinki. Subjects (or their parents or guardians) have given their written informed consent and that the study protocol was approved by the institute's committee on human research.

Author Contributions

All authors contributed to data analysis, drafting or revising the article, gave final approval of the version to be published, agreed to the submitted journal, and agree to be accountable for all aspects of the work.

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Disclosure

The authors have no conflicts of interest to declare.

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