

ROLE OF FIRST TRIMESTER HbA1c AS A PREDICTOR OF ADVERSE OBSTETRIC OUTCOMES IN A MULTI-ETHNIC COHORT

L. Mañé¹, J.A. Flores-Le Roux^{1,2}, D. Benaiges^{1,2}, M. Rodríguez², I. Marcelo², J.J. Chillarón^{1,2}, G. Llauradó¹, J. Pedro-Botet^{1,2}, R. Carreras^{2,3}, A. Payà^{2,3}.

¹ENDOCRINOLOGY, Hospital Del Mar, Barcelona, Spain. ²DEPARTMENT OF MEDICINE. UAB university, Barcelona, Spain. ³GINECOLOGY, Hospital Del Mar, Barcelona, Spain.

Background. Risk of obstetric complications increases linearly with rising maternal glycemia. Testing HbA1c is an effective option to detect hyperglycemia but its association with adverse pregnancy outcomes remains unclear. Emerging data sustains that an early HbA1c $\geq 5.9\%$ could act as a pregnancy risk marker.

Aims. To determine, in a multi-ethnic cohort, the usefulness of a 5.9% first-trimester HbA1c threshold to identify women without diabetes at increased pregnancy risk.

Materials and methods. A prospective study was conducted between April 2013-September 2015. We included women over 18 years with a singleton pregnancy. All women had an HbA1c measurement added to their first antenatal bloods. Women diagnosed of unknown type 2 diabetes were referred to the Diabetes Unit while those with an HbA1c value $< 6.5\%$ did not undergo any additional testing until 24-28 weeks of gestation when they underwent routine gestational diabetes mellitus (GDM) screening using the two-step approach. **Primary outcome** was macrosomia. **Secondary outcomes** were pre-eclampsia, preterm birth and Caesarean section rate.

Results.

Table 1. Maternal and gestational characteristics and pregnancy outcomes stratified according to HbA1c measurement at 1st antenatal blood testing.

| | HbA1c < 5.9% n=1,180 n (%) | HbA1c 5.9-6.4% n=48 n (%) | P value |
|--|----------------------------------|---------------------------------|---------|
| Maternal and gestational characteristics | | | |
| Age, years, mean \pm SD | 32.61 \pm 5.69 | 33.70 \pm 5.232 | 0.194 |
| Pre-pregnancy BMI, Kg/m ² , mean \pm SD | 25.34 \pm 5.04 | 28.10 \pm 5.35 | 0.001 |
| Previous macrosoma | 32/1113 (2.9) | 2/45 (4.4) | 0.384 |
| Multiparous | 644/1170 (55) | 33/48 (68.7) | 0.075 |
| GDM diagnosis | 129/1109 (11.6) | 22/47 (46.8) | <0.001 |
| Ethnicity | | | |
| -Caucasian | 614/1132 (54.2) | 17/45 (37.8) | 0.006 |
| -South Central Asian | 202/1132 (17.8) | 18/45 (40.0) | |
| -Moroccan | 81/1132 (7.2) | 2/45 (4.4) | |
| -Latin American | 155/1132 (13.7) | 4/45 (8.9) | |
| -East Asian | 63/1132 (5.6) | 4/45 (8.9) | |
| -Other | 17/1132 (1.5) | 0 | |
| Pregnancy weight gain, Kg, mean \pm SD | 10.886 \pm 4.66 | 9.52 \pm 5.11 | 0.055 |
| Anemia | 78 (6.6) | 9 (18.8) | 0.001 |
| Microcytosis | 136 (11.5) | 16 (33.3) | <0.001 |
| Pregnancy outcomes | | | |
| Preeclampsia | 43/1115 (3.9) | 4/43 (9.32) | 0.092 |
| Preterm birth | 78/1155 (6.8) | 5/47 (10.6) | 0.369 |
| Cesarean section | 313/1169 (26.8) | 14/47 (29.8) | 0.312 |
| Macrosomia | 69/1171 (5.9) | 8/48 (16.7) | 0.008 |

Table 2. Maternal and gestational characteristics and pregnancy outcomes among women with HbA1c $\geq 5.9\%$ stratified according to GDM diagnosis.

| | NO GDM (n=25)n (%) | GDM (n=22) n (%) | P value |
|--|-----------------------|---------------------|---------|
| Maternal and gestational characteristics | | | |
| Age, years, mean \pm SD | 33.54 \pm 5.72 | 33.81 \pm 4.87 | 0.865 |
| Pre-pregnancy BMI, Kg/m ² , mean \pm SD | 26.20 \pm 4.59 | 30.41 \pm 5.46 | 0.008 |
| Previous macrosoma | 0 | 2/20 (10) | 0.201 |
| Multiparous | 15/25 (60) | 14/20 (70) | 0.544 |
| Ethnicity | | | |
| -Caucasian | 12/24 (50) | 4/20 (20) | 0.020 |
| -South-Central Asian | 7/24 (29.9) | 11/20 (55) | |
| -Moroccan | 2/24 (8.3) | 0 | |
| -Latin American | 3/24 (12.5) | 1/20 (5) | |
| -East Asian | 0 | 4/20 (20) | |
| Pregnancy weight gain, Kg, mean \pm SD | 11.33 \pm 5.52 | 7.37 \pm 3.63 | 0.007 |
| Anemia | 7 (28) | 2 (9.1) | 0.062 |
| Microcytosis | 10 (40) | 6 (27.2) | 0.358 |
| Pregnancy outcomes | | | |
| Preeclampsia | 3/24 (12.5) | 1/18 (5.5) | 0.623 |
| Preterm birth | 2/25 (8.0) | 3/22 (13.6) | 0.654 |
| Cesarean section | 8/25 (32.0) | 5/21 (23.8) | 0.822 |
| Macrosomia | 5/25 (20) | 3/22 (13.6) | 0.706 |

Figure 1. Flow chart of the study protocol

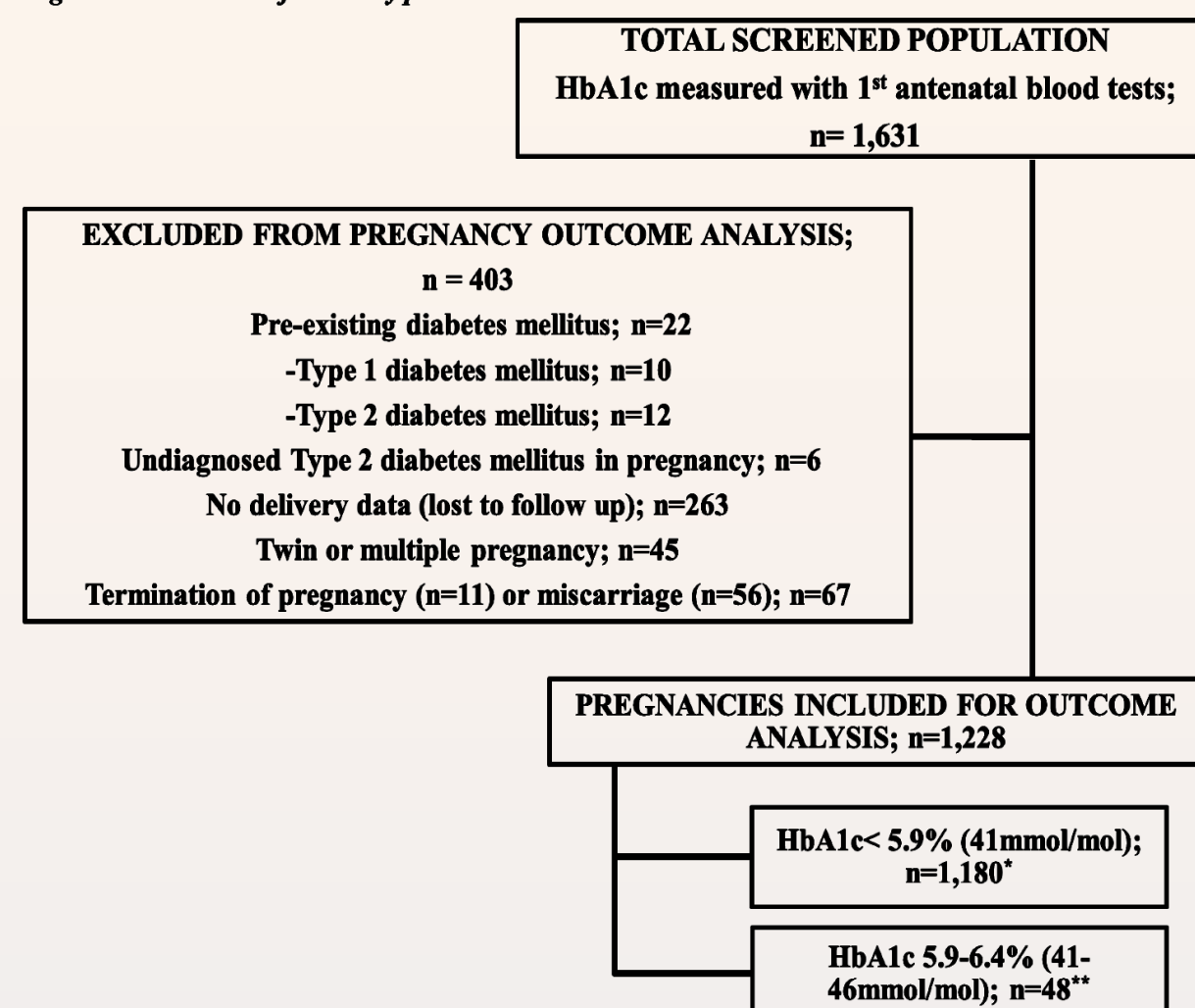
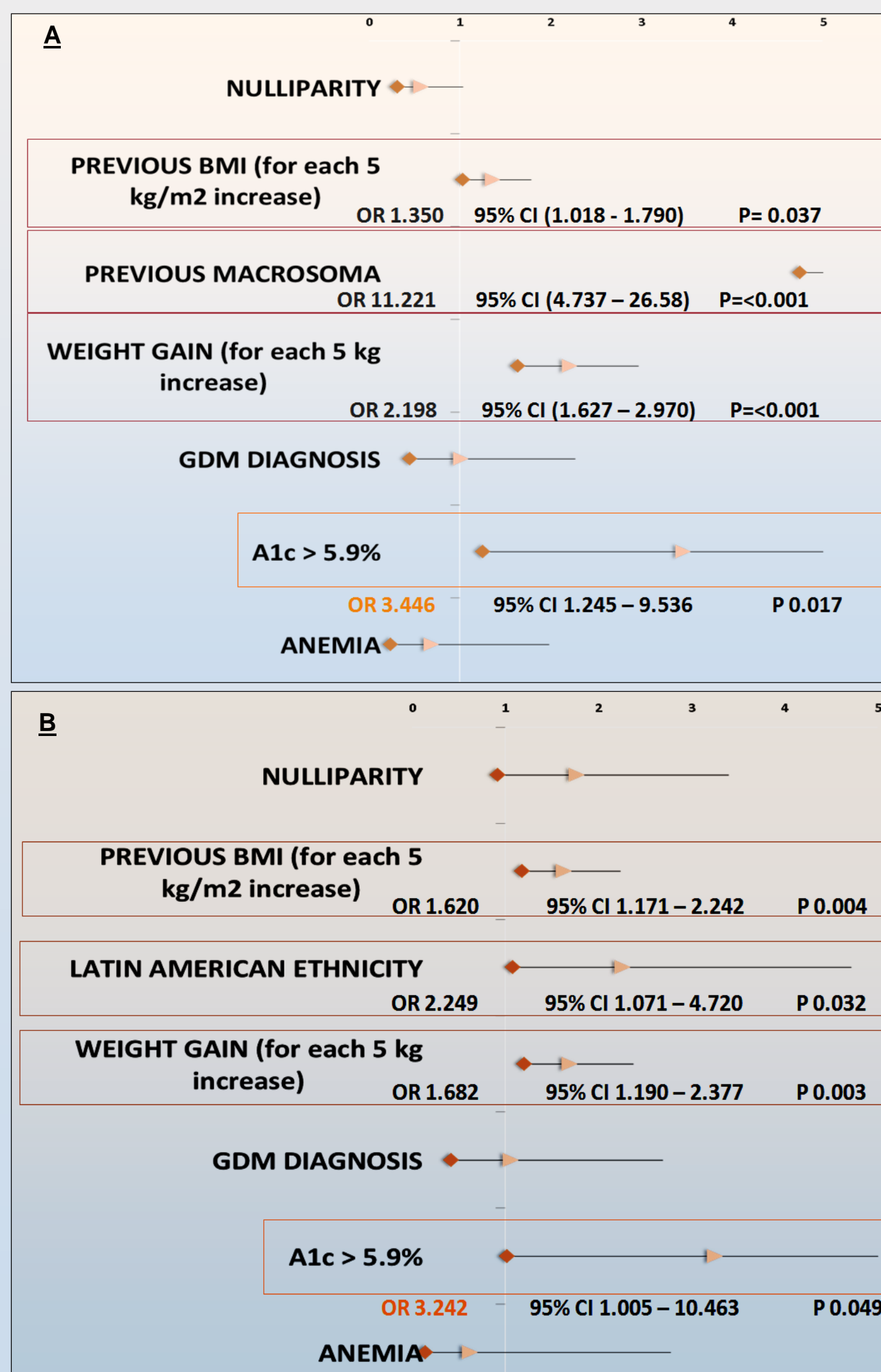


Fig 2. Multivariate analysis of predictor factors for macrosomia (A) and preeclampsia (B).



Conclusions. In a multiethnic population, a cutoff point of HbA1c $\geq 5.9\%$ in early pregnancy identifies a group of women at high risk for poorer pregnancy outcomes independently of GDM diagnosis later in pregnancy. Further studies are required to establish cutoff points adapted to each ethnic group and to assess whether early detection and treatment are of benefit.