Comparing IADPSG and NICE Diagnostic Criteria for GDM in Predicting Adverse Pregnancy Outcomes

OBJECTIVE
To compare the performance of diagnostic criteria for gestational diabetes mellitus (GDM) proposed by the International Association of the Diabetes and Pregnancy Study Groups (IADPSG) with those endorsed by the National Institute for Health and Care Excellence (NICE) in predicting adverse pregnancy outcomes.

RESEARCH DESIGN AND METHODS
We performed a secondary data analysis of the Hyperglycemia and Adverse Pregnancy Outcomes (HAPO) study participants in five study centers. Logistic regression analyses were performed, and Akaike information criterion were applied for the comparison of different statistical prediction models. We further analyzed the performance by four racial/ethnic subgroups, namely, Whites, Hispanics, Asians, and Blacks.

RESULTS
Among all, IADPSG criteria diagnosed 267 (4.1%) more women with GDM, but predicted primary caesarean section (CS) and large for gestational age (LGA) and neonatal adiposity better than did NICE criteria after adjustment for potential confounders. Among Whites, IADPSG criteria diagnosed 65 (2.5%) more subjects with GDM and predicted LGA and neonatal adiposity better, but predicted hypertensive disorders, primary CS and clinical neonatal hypoglycemia worse. Among Hispanics, the IADPSG criteria diagnosed 203 (12.1%) more with GDM but performed better in predicting hypertensive disorders, LGA, neonatal adiposity, and hyperinsulinemia. Among Asians, the IADPSG criteria diagnosed 34 (2.0%) fewer subjects with GDM but predicted hypertensive disorders better in the unadjusted model. In Blacks, IADPSG criteria diagnosed 34 (10.5%) more women with GDM.

CONCLUSIONS
IADPSG criteria appear to be more favorable than NICE for identification of adverse pregnancy outcomes among Hispanic and Asian women, while they are comparable to NICE among White women.
However, they have not yet been universally adopted. In 2015, the National Institute for Health and Care Excellence (NICE) of the U.K. developed new diagnostic criteria for GDM based on a health economic modeling for immediate pregnancy complications. NICE (2015) proposed that a fasting plasma glucose (FPG) of 5.6 mmol/L (100 mg/dL) and a 2-h plasma glucose (PG) of 7.8 mmol/L (140 mg/dL) were more cost-effective compared with the IADPSG criteria based on the quality-adjusted life-year (QALY) and incremental cost-effectiveness ratio (ICER) (4). IADPSG criteria differ from those of NICE, with a substantially lower fasting glucose (5.1 mmol/L or 92 mg/dL) and a higher 2-h glucose value (8.5 mmol/L or 153 mg/dL), with an additional 1-h glucose level (10.0 mmol/L or 180 mg/dL).

Several studies have compared the performance of both criteria in predicting adverse pregnancy outcomes, but the conclusions are not uniform. The main limitations of previous studies were that the studies were mostly confined to Caucasian populations and that their treatment criteria varied.

In the original Hyperglycemia and Adverse Pregnancy Outcome (HAPO) study, on which the IADPSG criteria were based (5), there was heterogeneity in the frequency of abnormal FPG, 1-h PG, and 2-h PG in women diagnosed with GDM across the 15 study centers (6). For example, an abnormal FPG level occurred in only 26% of women in the Hong Kong center but in >70% of women in Bellflower (7,8). Although FIGO advocated universal adoption of IADPSG criteria for the diagnosis of GDM, it remains uncertain whether the IADPSG criteria are superior to other existing diagnostic criteria in all ethnicities.

Because in clinical practice diagnosis of GDM is followed by treatment, it is extremely difficult to compare the application of different diagnostic criteria to the same patients who were treated for GDM by one set of criteria while they may or may not have been treated under another set of criteria. Furthermore, several studies have reported that women who met the IADPSG criteria but were diagnosed as non-GDM based on the NICE criteria had significantly increased risks of large for gestational age (LGA) infants and caesarean sections (CS) (9–12). Nevertheless, the diagnostic criteria in those studies were either heterogenous or unknown. In the Cambridge historical cohort, the diagnostic criteria adopted for GDM management were WHO 1999 or modified WHO 1999, which were quite similar to the NICE 2015 criteria (10). A study of the Finnish population was, however, based on the local criteria (i.e., FPG 5.3 mmol/L, 1-h 10.0 mmol/L, 2-h 8.6 mmol/L) (11). Lastly, a study of the Indian population did not specify the criteria used for GDM treatment, so it is difficult to have a head-to-head comparison of the performance between the two criteria (12).

Our aim was to use a subset of the HAPO study data obtained from five study centers where all participants received no intervention for GDM and where both the clinicians and participants were blinded from the oral glucose tolerance test (OGTT) results.

**RESEARCH DESIGN AND METHODS**

**Study Design**

The original HAPO study protocol has been described previously elsewhere (13). We included data from five HAPO study centers: 1) Bellflower, CA; 2) Cleveland, OH; 3) Brisbane, Queensland, Australia; 4) Newcastle, New South Wales, Australia; and 5) Hong Kong, China.

All pregnant women booked for antenatal care in the study centers were eligible to participate unless they had one or more of the following exclusion criteria: age <18 years, a plan to undergo tropin ovulation induction or in vitro fertilization, glucose testing before recruitment or a diagnosis of diabetes during the current pregnancy, diagnosis of diabetes before the current pregnancy and requiring treatment with medication, participation in another study that could interfere with the HAPO study, or infection with HIV or hepatitis B or C virus.

All participants underwent a 75-g OGTT at 24–32 weeks of gestation. Women with a FPG level of >105 mg/dL (5.8 mmol/L) or 2-h plasma glucose level of >200 mg/dL (11.1 mmol/L) were unblinded and received treatment for GDM accordingly. Otherwise, participants and health care providers were blinded to the OGTT results, and participants received no treatment. All maternal and umbilical cord blood samples were analyzed at the central laboratory of the HAPO study.

Self-reported race/ethnicity (i.e., White, Hispanic, Asian [majority Chinese], Black, and others/miscellaneous), maternal age, gestational age at delivery, BMI at OGTT, smoking status, and parity (nulliparity vs. multiparity) were extracted as demographic and clinical characteristics. Neonatal measurements were obtained by a standardized method within 72 h after the delivery. Infants’ percentage of body fat was estimated as $(0.39055 \text{ [birth weight]} + 0.0453 [\text{flank skinfold}] - 0.03237 [\text{length}] + 0.54657)/[\text{birth weight}])$, as previously described (14).

**Diagnostic Definition of GDM by NICE and IADPSG Criteria**

Women whose FPG was ≥5.6 mmol/L (100 mg/dL) and/or 2-h PG was ≥7.8 mmol/L (140 mg/dL) were defined as GDM by NICE criteria, while those whose FPG was ≥5.1 mmol/L (92 mg/dL) and/or 1-h PG was ≥10.0 mmol/L (180 mg/dL) and/or 2-h PG was ≥8.5 mmol/L (153 mg/dL) were defined as GDM by IADPSG criteria. Furthermore, all women were classified under one of the four categories: 1) non-GDM by both IADPSG and NICE; 2) GDM by IADPSG but non-GDM by NICE; 3) GDM by NICE but non-GDM by IADPSG; and 4) GDM by both the IADPSG and NICE criteria.

**Outcome Variables**

We compared participants who remained blinded on six adverse pregnancy outcomes: 1) hypertensive disorder of pregnancy, namely, gestational hypertension (GH) or preeclampsia (PE); 2) primary CS; 3) LGA infants (infants born with birth weight >90th percentile adjusted for ethnicity, study center, gestational age, parity, and infant’s sex); 4) neonatal adiposity (percentage body fat >90th percentile adjusted for ethnicity, study center, gestational age, parity, and infant’s sex); 5) clinical neonatal hyperglycemia (being present if there was a notation of neonatal hyperglycemia in the medical record and there were symptoms or treatment with a glucose infusion or a local laboratory report of a glucose value of ≤1.7 mmol/L in the first 24 h after birth or ≤2.5 mmol/L after the first 24 h [13]); and 6) neonatal hyperinsulinemia (umbilical cord blood
C-peptide level >90th percentile; i.e., 1.7 μg/L (15,16), among all blinded participants and among four ethnic subgroups (i.e., Whites, Hispanics, Asians, and Blacks).

Statistical Analyses
Continuous variables are expressed as mean ± SD and were compared by using the Kruskal-Wallis test. Post hoc pairwise analyses were corrected by the Wilcoxon test when the null hypotheses were rejected. Categorical variables are expressed as n (%) and were compared by the McNemar test, χ² test, or Fisher exact test, as appropriate, for between-group comparisons and specified in legends of the tables. If the null hypothesis was rejected, post hoc pairwise comparisons were further conducted, corrected by the false discovery rate. Logistic regressions were used to explore the association of GDM by IADPSG or NICE criteria with adverse pregnancy outcomes. Data were adjusted for maternal age, parity, race/ethnicity, and study center (model 1), and model 1 plus maternal height, BMI and gestational age at OGTT, smoking status, alcohol use, maternal urinary tract infection (for GH/PE only), hospitalization before delivery, and mean arterial pressure at OGTT (except for GH/PE), family history of diabetes, and baby’s sex (model 2) (13). We also applied Akaike information criterion (AIC) for the comparison of different statistical prediction models in the logistic regression analysis (17). A model with a lower AIC value of >4 was considered as significantly better in performance (17). P values <0.05 were used to indicate significance for two-tailed statistical test results. All statistical analyses were performed by using R 4.0.3 software (downloadable at www.r-project.org).

RESULTS
Among 6,544 women recruited from the five HAPO study centers, 1,223 (18.7%) and 956 women (14.6%) were diagnosed with GDM by the IADPSG and NICE criteria, respectively (Table 1). Among the 1,732 participants in the Hong Kong center, 1,595 were Chinese, suggesting at least 92% of the Asians in this cohort were of Chinese race/ethnicity. IADPSG diagnosed significantly more women with GDM than did NICE criteria among Whites (15.7% vs. 13.2%, P < 0.001), Hispanics (26.4% vs. 14.3%, P < 0.001), and Blacks (20.7% vs. 10.2%, P < 0.001), but less GDM among Asians (15.7% vs. 17.7%, P = 0.011). When comparing by study centers, IADPSG diagnosed more women with GDM than did NICE among women from the two U.S. centers (i.e., Bellflower and Cleveland), but less GDM among women from the Hong Kong center. (Table 1). There were no differences in the incidence of GDM by the two diagnostic criteria in either of the Australian centers. In this multiracial and multiethnic cohort, Hispanic women have the highest GDM rate if IADPSG criteria were adopted, while Asian women (majority Chinese) have the highest rate if NICE diagnostic criteria were used (Supplementary Fig. 1); whereas, the frequency of women being non-GDM by both criteria was lowest in Hispanics but highest in Whites. In contrast, the frequency of women with GDM by both criteria was highest in Hispanics but lowest in Whites (Supplementary Table 1).

Table 1—The frequencies of GDM diagnosed according to IADPSG and NICE criteria by races/ethnicities and HAPO study centers

| Race/ethnicities | Participants (n) | IADPSG (%) | NICE (%) | P        |
|------------------|------------------|------------|----------|----------|
| Overall          | 6,544 (100)      | 1,223 (18.7) | 956 (14.6) | <0.001   |
| **Race**         |                  |            |          |          |
| White            | 2,583 (39.5)     | 406 (15.7) | 341 (13.2) | <0.001   |
| Hispanic         | 1,677 (25.6)     | 442 (26.4) | 239 (14.3) | <0.001   |
| Asian*           | 1,732 (26.5)     | 272 (15.7) | 306 (17.7) | 0.011    |
| Black            | 324 (5.0)        | 67 (20.7)  | 33 (10.2)  | <0.001   |
| Other            | 228 (3.5)        | 36 (15.8)  | 37 (16.2)  | 0.835    |
| **Centers**      |                  |            |          |          |
| Bellflower       | 1,981 (30.3)     | 505 (25.5) | 271 (13.7) | <0.001   |
| Cleveland        | 797 (12.2)       | 199 (25.0) | 134 (16.8) | <0.001   |
| Brisbane         | 1,444 (22.1)     | 179 (12.4) | 171 (11.8) | 0.505    |
| Newcastle        | 668 (10.2)       | 102 (15.3) | 93 (13.9)  | 0.292    |
| Hong Kong        | 1,654 (25.3)     | 238 (14.4) | 287 (17.4) | <0.001   |

Data are expressed as n (%). Frequencies were compared by the McNemar test. *More than 92% of the Asians in the cohort were Chinese ethnicity.
Supplementary Table 4 compares the performance of IADPSG with that of NICE criteria in the prediction of maternal and neonatal adverse outcomes among all women whose OGTT result remained blinded. The IADPSG performed better than NICE criteria in predicting hypertensive disorders, primary CS, LGA infants, higher neonatal adiposity, and hyperinsulinemia, indicated by AIC, after adjustment for maternal age, parity, race/ethnicity, and study center (model 1). However, the performance of both IADPSG and NICE were attenuated with further adjustment for confounders in model 2, which weakened the better performance of IADPSG in predicting hypertensive disorders and neonatal hyperinsulinemia (model 2). Clinical neonatal hypoglycemia was only predicted by the IADPSG, but not the NICE criteria in the unadjusted regression model. However, the performances of the IADPSG and NICE criteria were attenuated and enhanced, respectively, after adjustment for the potential confounders (model 1 and 2), which in contrast, made the clinical neonatal hypoglycemia only predicted by the NICE but not the IADPSG criteria (Supplementary Table 4).

Table 2 shows the result of comparisons in Whites. IADPSG criteria performed better than NICE criteria in predicting LGA infants, indicated by AIC, after adjustment for maternal age, parity, race/ethnicity, and study center (model 1). However, the performance of both IADPSG and NICE were attenuated with further adjustment for confounders in model 2, which weakened the better performance of IADPSG in predicting hypertensive disorders and neonatal hyperinsulinemia (model 2). Clinical neonatal hypoglycemia was only predicted by the IADPSG, but not the NICE criteria in the unadjusted regression model. However, the performances of the IADPSG and NICE criteria were attenuated and enhanced, respectively, after adjustment for the potential confounders (model 1 and 2), which in contrast, made the clinical neonatal hypoglycemia only predicted by the NICE but not the IADPSG criteria (Supplementary Table 4).

Figure 1—The frequency distribution of plasma glucose levels at OGTT among the White (A, B, and C), Hispanic (D, E, and F), Asian (>92% of the Asians in the cohort were Chinese ethnicity) (G, H, and I), and Black (J, K, and L) women in the HAPO study subcohort. The dashed lines in orange and purple indicate the thresholds of the IADPSG and NICE criteria, respectively.
Table 2—Associations of GDM diagnosed by the IADPSG and NICE criteria with adverse pregnancy outcomes in Whites, whose OGTT remained blinded (n = 2,551)

| Outcome                  | Model 1 | Model 2 |
|--------------------------|---------|---------|
|                          | IADPSG  | NICE    |
|                          | (95% CI) | (95% CI) |
|                          | AIC     | AIC     |
| Maternal adverse outcomes|         |         |
| GH/PE                    | 1.99 (1.52–2.62) | 2.02 (1.52–2.62) |
| Primary CS               | 1.51 (1.03–2.16)  | 1.61 (1.06–2.39)  |
| Neonatal outcomes        |         |         |
| GH/PE                    | 2.11 (1.52–2.96)  | 2.20 (1.51–2.96)  |
| Adiposity                | 1.76 (1.09–2.95)  | 2.28 (1.40–3.70)  |
| Hyperinsulinemia         | 2.39 (1.67–3.39)  | 2.53 (1.73–3.66)  |
| Clinical hypoglycemia    | 1.92 (1.32–2.74)  | 2.14 (1.50–3.10)  |
| Hypertension             | 1.36 (0.94–1.96)  | 1.37 (0.94–1.96)  |

*P < 0.05; ‡P < 0.005; §P < 0.001

Our study compared the prevalence of GDM and performance in predicting pregnancy outcomes between the IADPSG and NICE criteria by using a subcohort of the HAPO study cohort and incorporating subgroup analysis by race/ethnicity (i.e., Whites, Hispanics, Asians [majority Chinese], and Blacks). The HAPO database provided a unique opportunity to examine adverse pregnancy outcomes independent of any intervention. Consistent with previous studies, we found that IADPSG criteria not only diagnosed more women as having GDM but also better predicted hypertensive disorders in pregnancy, primary CS, and LGA infants with LGA infants, after adjustment for confounders in model 2. Moreover, the IADPSG criteria also performed better in predicting LGA infants indicated by the AIC. Meanwhile, primary CS could only be predicted by IADPSG, but not NICE criteria, and confined to unadjusted model only (Table 3).

In Blacks, neonatal adiposity was only predicted by the IADPSG criteria, whereas primary CS and neonatal hyperinsulinemia were only predicted by the NICE criteria, after adjustment for confounders in model 2. On the other hand, the IADPSG criteria also performed better in predicting LGA infants indicated by the AIC value, after adjustment for confounders in model 2 (Table 4).

**CONCLUSIONS**

Our study compared the prevalence of GDM and performance in predicting pregnancy outcomes between the IADPSG and NICE criteria by using a subcohort of the HAPO study cohort and incorporating subgroup analysis by race/ethnicity (i.e., Whites, Hispanics, Asians [majority Chinese], and Blacks). The HAPO database provided a unique opportunity to examine adverse pregnancy outcomes independent of any intervention. Consistent with previous studies, we found that IADPSG criteria not only diagnosed more women as having GDM but also better predicted hypertensive disorders in pregnancy, primary CS, and LGA infants (10–12,18). In addition, we also found that IADPSG criteria performed better in detecting higher neonatal adiposity, hyperinsulinemia, and clinical hypoglycemia only by the IADPSG, whereas hypertensive disorders, primary CS, and clinical neonatal hypoglycemia were predicted only by the NICE criteria, based on the P value (Table 2).
### Table 3—Associations of GDM diagnosed by the IADPSG and NICE criteria with adverse pregnancy outcomes in Hispanics, whose OGTT remained blinded (n = 1,611)

|                      | Unadjusted | Adjusted                                      |
|----------------------|------------|----------------------------------------------|
|                      | IADPSG     | NICE                                        |
|                      | OR (95% CI)| AIC                                         |
|                      | OR (95% CI)| AIC                                         |
| Maternal adverse outcomes |           |                                              |
| GH/PE                | 1.72 (1.32–2.24) | 1,662.0                                    |
|                      | 1.41 (0.99–2.02) | 1,674.1                                    |
|                      | 1.81 (1.37–2.39) | 1,606.9                                    |
|                      | 1.49 (1.03–2.16) | 1,619.9                                    |
|                      | 1.43 (1.07–1.92) | 1,523.2                                    |
|                      | 1.43 (0.97–2.10) | 1,525.8                                    |
| Primary CS           | 1.74 (1.22–2.47) | 1,011.3                                    |
|                      | 1.38 (0.84–2.19) | 1,018.7                                    |
|                      | 1.45 (0.99–2.10) | 917.7                                      |
|                      | 1.07 (0.62–1.76) | 921.2                                      |
|                      | 1.29 (0.86–1.89) | 907.3                                      |
|                      | 0.96 (0.55–1.59) | 908.8                                      |
| Neonal outcomes      |            |                                              |
| LGA                  | 2.33 (1.64–3.29) | 989.2                                      |
|                      | 1.78 (1.11–2.77) | 1,004.8                                    |
|                      | 2.35 (1.64–3.35) | 993.9                                      |
|                      | 1.77 (1.10–2.78) | 1,009.4                                    |
|                      | 2.01 (1.38–2.90) | 971.1                                      |
|                      | 1.88 (1.14–2.99) | 977.9                                      |
| Adiposity            | 1.94 (1.29–2.89) | 780.3                                      |
|                      | 1.13 (0.60–1.98) | 789.8                                      |
|                      | 2.04 (1.34–3.07) | 784.2                                      |
|                      | 1.18 (0.62–2.08) | 794.7                                      |
|                      | 1.62 (1.04–2.50) | 765.8                                      |
|                      | 1.20 (0.62–2.16) | 770.0                                      |
| Clinical hyperglycemia | 1.09 (0.05–8.52) | 59.9                                       |
|                      | —          | —                                            |
|                      | 0.95 (0.05–7.98) | 61.0                                      |
|                      | —          | —                                            |
|                      | 0.79 (0.04–7.22) | 71.0                                      |
|                      | —          | —                                            |
| Hyperinsulinemia      | 2.34 (1.63–3.33) | 927.8                                      |
|                      | 1.81 (1.12–2.85) | 942.6                                      |
|                      | 2.18 (1.51–3.14) | 927.3                                      |
|                      | 1.65 (1.01–2.62) | 939.9                                      |
|                      | 1.98 (1.35–2.90) | 922.5                                      |
|                      | 1.52 (0.92–2.43) | 931.7                                      |

**Model 1 was adjusted for maternal age, parity, ethnicity, and study center. Model 2 was adjusted for covariates in model 1 plus maternal height, BMI, and gestational age at OGTT, smoking status, alcohol use, maternal urinary tract infection (for GH/PE only), hospitalization before delivery, and mean arterial pressure at OGTT (except for GH/PE), family history of diabetes, and baby’s sex. OR, odds ratio; — inadequate number of patients for analysis. *P < 0.05; †P < 0.01; ‡P < 0.005; §P < 0.001.**
### Table 4: Associations of GDM diagnosed by the IADPSG and NICE criteria with adverse pregnancy outcomes in Asians, whose OGTT remained blinded (n = 1,695)

|                        | Model 1 | Model 2 |
|------------------------|---------|---------|
|                        | IADPSG  | NICE    |
|                        | OR      | (95% CI)| AIC     | OR      | (95% CI)| AIC     |
| Maternal adverse outcomes |        |         |         |        |         |         |
| GH/PE                  | 1.60 (1.06–2.41)* | 1.079.2 | 1.12 (0.73–1.71) | 1.031.5 | 1.099.6 (0.63–1.60) | 1.003.8 |
| Primary CS             | 1.36 (0.88–2.10) | 1.029.7 | 1.04 (0.66–1.61) | 1.031.5 | 1.036.0 (0.69–1.69) | 1.003.8 |
|                        | 1.573.8 | 1.06 (0.72–1.53) | 1.031.5 | 1.048.0 (0.69–1.39) | 1.003.8 |
|                        |         |         |         |        |         |         |
| Adjusted Model 1 Model 2 |         |         |         |        |         |         |
|                        | IADPSG  | NICE    |
|                        | OR      | (95% CI)| AIC     | OR      | (95% CI)| AIC     |
| Maternal adverse outcomes |        |         |         |        |         |         |
| GH/PE                  | 1.60 (1.06–2.41)* | 1.079.2 | 1.12 (0.73–1.71) | 1.031.5 | 1.099.6 (0.63–1.60) | 1.003.8 |
| Primary CS             | 1.36 (0.88–2.10) | 1.029.7 | 1.04 (0.66–1.61) | 1.031.5 | 1.036.0 (0.69–1.69) | 1.003.8 |
|                        | 1.573.8 | 1.06 (0.72–1.53) | 1.031.5 | 1.048.0 (0.69–1.39) | 1.003.8 |
|                        |         |         |         |        |         |         |
| Neonatal outcomes      |        |         |         |        |         |         |
| GH/PE                  | 1.80 (1.19–2.52) § | 1.059.2 | 1.70 (1.11–2.54) § | 1.060.4 | 1.63 (1.09–2.39)* | 1.060.8 |
| Primary CS             | 1.54 (0.97–2.46) § | 1.04 (0.69–1.62) § | 1.04 (0.66–1.61) | 1.031.5 | 1.048.0 (0.69–1.39) | 1.003.8 |
|                        | 1.87 (1.12–2.61) § | 1.057.5 | 1.71 (1.10–2.46) § | 1.058.0 | 1.63 (1.09–2.39)* | 1.060.8 |
|                        |         |         |         |        |         |         |
| Adjusted Model 1 Model 2 |         |         |         |        |         |         |
|                        | IADPSG  | NICE    |
|                        | OR      | (95% CI)| AIC     | OR      | (95% CI)| AIC     |
| Neonatal outcomes      |        |         |         |        |         |         |
| GH/PE                  | 1.80 (1.19–2.52) § | 1.059.2 | 1.70 (1.11–2.54) § | 1.060.4 | 1.63 (1.09–2.39)* | 1.060.8 |
| Primary CS             | 1.54 (0.97–2.46) § | 1.04 (0.69–1.62) § | 1.04 (0.66–1.61) | 1.031.5 | 1.048.0 (0.69–1.39) | 1.003.8 |
|                        | 1.87 (1.12–2.61) § | 1.057.5 | 1.71 (1.10–2.46) § | 1.058.0 | 1.63 (1.09–2.39)* | 1.060.8 |
|                        |         |         |         |        |         |         |

Model 1 was adjusted for maternal age, parity, ethnicity, and study center. Model 2 was adjusted for covariates in model 1 plus maternal height, BMI, and gestational age at OGTT, smoking status, alcohol use, family history of diabetes, and baby’s sex. *P < 0.05; †P < 0.01; §P < 0.001; $P < 0.0001.

More than 92% of the Asians in the cohort were Chinese ethnicity.

Lastly, IADPSG criteria diagnosed fewer women as having GDM among Asians, but better predicted women with hypertensive disorders in pregnancy, limited to the unadjusted model. Therefore, our results support the use of IADPSG criteria in an Asian population. However, most of the Asians in our cohort were Chinese; whether this result is generalizable to other Asian populations, such as South Asians, is uncertain.

Our results also illustrate the underlying reason for the difference in the prevalence of GDM between IADPSG and NICE criteria in Asians (majority Chinese, Hispanics, and Blacks. The difference lies in the proportion of women whose FPG levels fell between 5.1 and 5.6 mmol/L, 1-h PG levels ≥10.0 mmol/L, and the 2-h PG levels between 7.8 and 8.5 mmol/L. The former two contribute to a higher GDM frequency using IADPSG criteria, while the third contributes to the extra GDM by the NICE criteria. Since fewer Asians in the cohort fell into the range of FPG level between 5.1 and 5.6 mmol/L, but more fell into a 2-h PG level between 7.8 and 8.5 mmol/L, IADPSG diagnosed 2% less GDM than NICE criteria in this population. In contrast, Hispanic women more often fell into the range of FPG levels between 5.1 and 5.6 mmol/L and 1-h PG level ≥10.0 mmol/L, but less often into the range of 2-h PG levels between 7.8 and 8.5 mmol/L, resulting in a 12.1% higher rate of GDM with IADPSG than with NICE criteria in Hispanics. Similarly, a 10.5% higher rate of GDM with IADPSG than NICE criteria in Blacks was attributed to more women falling into the FPG level range between 5.1 and 5.6 mmol/L, but fewer falling into the 2-h PG level range between 7.8 and 8.5 mmol/L.

The current study has several strengths over previous studies comparing the IADPSG and NICE diagnostic criteria. Firstly, none of our participants received any treatment for hyperglycemia, and neither the women nor the managing clinicians were made aware of the OGTT results, in contrast to heterogeneous treatment criteria among the previous studies. The performance of NICE criteria in the Cambridge cohort might have been
underestimated because more women being diagnosed with GDM by the NICE criteria were treated accordingly instead of those diagnosed with the IADPSG criteria (10).

Secondly, our study constituted a multi-ethnic population that enabled subgroup comparisons by races/ethnicities. The review by the NICE 2015 committee as well as in other previous studies was predominantly in Whites.

Thirdly, the original HAPO study design enabled us to compare additional outcomes, including neonatal adiposity and hyperinsulinemia, which were found highly relevant to long-term metabolic health in recent studies (21,22).

Lastly, our data set also allowed the comparison of both sets of criteria after adjusting for multiple potential confounders associated with GDM and other adverse pregnancy outcomes, such as family history of diabetes and maternal BMI. Maternal BMI at midgestation is contributed to by prepregnant BMI and gestational weight gain; both maternal obesity and excessive gestational weight gain are independent risk factors for hypertensive disorders, primary CS, LGA, and neonatal adiposity (23–26).

Our study has also some limitations. For example, our data set did not have certain outcomes known to be associated with maternal hyperglycemia, namely, spontaneous preterm prelabor rupture of membranes, shoulder dystocia, neonatal jaundice, or respiratory distress syndrome.

Secondly, the Asian subgroup was mainly confined to the Chinese from the Hong Kong center; the application of IADPSG and NICE criteria to other Asian groups, such as South Asian, might be different.

Thirdly, our study did not have the information to compare the cost-effectiveness between the two sets of criteria. Nevertheless, IADPSG criteria apparently can identify fewer women as having GDM but can identify more adverse outcomes among those so identified in the Chinese population.

Lastly, HAPO study was an observational study. We still need a randomized study to address whether treatment of GDM based on IADPSG versus NICE criteria offers better pregnancy outcomes, as was suggested previously (27).

In this multiracial/ethnic population, the IADPSG criteria diagnosed more Hispanics with GDM than the NICE criteria and performed better in the prediction of four individual adverse pregnancy outcomes in this subgroup. However, among Asians, the majority of whom were Chinese ethnicity, the IADPSG criteria diagnosed fewer women with GDM, while predicting hypertensive disorders better. Among Whites, the IADPSG criteria diagnose more women with GDM. Nonetheless, either IADPSG or NICE criteria performed better with relation to prediction of specific outcomes. One of the original goals of the HAPO study was to develop uniform standard criteria for the diagnosis of GDM using a 75-g OGTT. These criteria could then be applied worldwide based on the large number of study subjects from diverse racial/ethnic groups in order to compare incidence rates and outcomes based on various treatment strategies. Our results indicate that differences in rates and type of adverse outcomes vary depending on whether IADPSG or NICE criteria are used in different racial/ethnic groups. These differences are most likely the result of population differences in maternal genetics, metabolism, nutrition, and activity. Understanding these differences will help caregivers and organizations make informed choices about which criteria would be best applied to an individual population. Based on our results, we suggest that each country determine the diagnostic criteria for GDM based on its racial/ethnic composition as well health economics.

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