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Evaluation of the Modified FINDRISC Diabetes Score to Identify Individuals at High Risk for Diabetes among Middle-aged White and Black ARIC Study Participants

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Abstract

Objective—To evaluate a modified Finnish Diabetes Risk Score (FINDRISC) score for predicting the risk of incident diabetes among white and black middle-aged Atherosclerosis Risk in Communities (ARIC) study participants.

Research Design and Methods—We assessed 9,754 ARIC cohort participants who were free of diabetes at baseline. Logistic regression and receiver operator characteristic (ROC) curves were used to evaluate a modified FINDRISC score for predicting incident diabetes after 9 years of

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DUALITY OF INTEREST

A.M.M. is an employee of, and owns stock in, Merck & Co. Inc.

AUTHORS CONTRIBUTIONS

M.M.K. assisted with data analysis, and wrote and edited the manuscript. C.G., S.H.G., W.D.R., B.D., and M.I.S. provided subject matter expertise and edited the manuscript. J.T. developed the FINDRISC and edited the manuscript. A.M.M. designed the study, oversaw the data collection, and wrote and edited the manuscript. R.E.F. conducted data analysis and wrote and edited the manuscript, and is the guarantor of this work and, as such, had full access to all the data in the study and takes responsibility for the integrity of the data and the accuracy of the data analysis.

Results—The mean FINDRISC score [range: 2 (lowest risk) -17 (highest risk)] for black women was higher (9.9±3.6) than for black men (7.6±3.9), white women (8.0±3.6), and white men (7.6±3.5). The incidence of diabetes generally increased across deciles of FINDRISC score for all four race/gender groups. ROC curve statistics for the FINDRISC score showed the highest area under the curve for white women (0.77) and lowest for black men (0.70).

Conclusions—We used a modified FINDRISC score to predict the 9-year risk of incident diabetes in a biracial United States population. The modified risk score can be useful for the early screening of incident diabetes in biracial populations, which may be helpful for early interventions to delay or prevent diabetes.

INTRODUCTION

The worldwide prevalence of diabetes was 8% in 2012 and approximately 5.1 million people aged between 20 and 79 years died from diabetes in 2013.¹ The prevalence of diabetes in the adult population of the United States (US) had exceeded 11% by the year 2010.² What many believe to be an epidemic of type 2 diabetes around the world has fueled interest in the development of simple and affordable screening strategies to identify participants who would benefit from aggressive lifestyle or pharmacologic prevention strategies.^{3,4}

To improve upon the predictive properties of fasting or 2-hour glucose tolerance, several multivariable models have been published⁵⁻¹³ that utilize a combination of risk factor profiles and measures of glucose disturbances with clinical variables. Although such "complex" algorithms may be appropriate for etiologic investigations of the underlying causes of the development of type 2 diabetes, they may be impractical from the standpoint of public health screening efforts to identify individuals at high-risk of glucose disturbances who would benefit principally from primary prevention strategies.^{5,6,9,10}

There are multiple reviews of diabetes risk scores in the literature.^{14–16} A recent database search yielded a total of 8,864 records for diabetes risk scores, but only 145 of them were actually tested in populations, and details of only 96 models were available.¹⁴ These studies took place in 17 countries on six different continents: 30 in Europe, 25 in North America, 21 in Asia, 8 in Australasia, 8 in the Middle East, 1 in South America, and 1 in Africa.¹⁵ Some of the simpler diabetes risk scores include Finnish Diabetes Risk Score (FINDRISC),¹⁷ ADA,¹⁸ Cambridge,¹⁹ German Risk Score,²⁰ Rotterdam,²¹ and Inter99²² which were developed to screen for prevalent or incident diabetes, based on a combination of age, family history of diabetes, history of gestational diabetes, anthropometric measures (weight, height, body mass index [BMI], waist circumference), levels of physical activity, dietary habits, use of anti-hypertensives or corticosteroids, and other constructs that do not require laboratory measurements. In Europe, the FINDRISC has been recommended by the European Society of Cardiology (ESC) and European Association of the Study of Diabetes (EASD) for the general population.²³

Specifically, the FINDRISC¹⁷ has been externally validated to identify participants at high risk for incident diabetes and to predict prevalent undiagnosed diabetes, impaired fasting glucose, or the metabolic syndrome in Finnish, Italian, and Greek populations.^{17, 24} Recently, a cross-sectional analysis of the FINDRISC risk score was carried out with NHANES data to identify participants with undiagnosed diabetes and pre-diabetes in US populations.²⁵ The purpose of the current study was to prospectively evaluate the ability of the FINDRISC to predict diabetes (treated or untreated) during 9 years of follow up among white and black middle-aged men and women in the US.

METHODS

Data from baseline (Visit 1) of the Atherosclerosis Risk in Communities (ARIC) study was used to identify participants with prevalent diabetes and to ascertain participants' values on FINDRISC algorithm components. Follow-up visits 2, 3, and 4 (occurring at approximately 3-year intervals) were used to ascertain incident diabetes status. Since not all self-reported FINDRISC data were available for analysis, we evaluated the diagnostic properties of a modified FINDRISC using receiver operator characteristic (ROC) curves. We assessed the ability of the modified FINDRISC to predict incident diabetes during 9 years of follow-up, for the overall population and within race/gender groups, in participants with no evidence of diabetes at baseline.

Study population

The ARIC study is a prospective study of 15,792 people aged 45–64 years from four US communities:^{26–27} Forsyth County, North Carolina, Jackson, Mississippi, suburbs of Minneapolis, Minnesota, and Washington County, Maryland. We excluded 48 participants who were not white or black, and blacks living in the Minnesota and Maryland study communities (n=55), due to a lack of power to make inferences to these populations. We also excluded 516 participants at baseline who were not fasting 8 hours or more, as well as 71 participants with unknown fasting status at the time of the baseline exam. An additional 1,611 participants were excluded for having diabetes at baseline and 95 were not included due to missing prevalent diabetes status at baseline. The number of participants missing data on FINDRISC score components such as BMI (kg/m²), waist circumference (cm), hypertensive medication use within the last two weeks, and family history of diabetes (paternal and maternal, living, or deceased) was 1,279. Finally, 2,363 participants were excluded for not having complete information on incident diabetes status throughout 9 year follow-up, resulting in 9,754 participants available for this analysis.

Study definition of diabetes

The definition of diabetes was the same for all four visits: a fasting blood glucose concentration of 126 mg/dl, self-reported, diagnosis by a physician, or the use of any medication for diabetes or high blood glucose.

Page 3

FINDRISC

The FINDRISC algorithm was developed in the Finnish population to screen for individuals at high risk of developing diabetes¹⁷ and includes the following measures: age; BMI; waist circumference; physical activity; daily consumption of fruits, vegetables, and berries; history of medication use for hypertension; self-reported history of elevated glucose (or gestational diabetes); and family history of diabetes (Table S1). We used a modified FINDRISC due to characteristics of our study population and available data, as described below. The comparison of the original FINDRISC with our modified FINDRISC with respect to variables and their associated score points is given in Table S1. The original¹⁷ FINDRISC ranged from 0–26 while our modified FINDRISC ranged from 2–17.

The age strata of eligible ARIC participants at baseline corresponded to the FINDRISC algorithm of 45–54 years and 55–64 years with two and three points, respectively. Meanwhile, the anthropometric measures of BMI and waist circumference were assessed at baseline for each cohort member, and defined exactly as shown in Table 1, consistent with the original FINDRISC.

Data on physical activity and fruit, vegetable, and berry consumption were not collected in ARIC at baseline in a manner consistent with the original¹⁷ FINDRISC (Table S1). Specifically, while participants were asked about physical activity at work and during leisure time, duration of activity was not captured. Similarly, data were ascertained at baseline regarding participants' consumption of particular fruits and vegetables, but not fruits, vegetables, and berries as food categories. Thus, we were not able to consider questions on physical activity and diet in the present analysis. It should be noted, however, that a simplified FINDRISC model without indicators of fruit and vegetable consumption and physical activity was successfully validated to predict type 2 diabetes risk prevalence in an Omani population²⁸ and with disease incidence without the dietary question in a German population.²⁹

Participants were asked to bring containers of all medications used in the past two weeks to the baseline visit. ARIC study staff additionally asked participants about specific types of medications, including, "*Were any of the medications you took during the past two weeks for high blood pressure?*" These data were used to ascertain whether a participant was taking high blood pressure medication on a regular basis (Table S1). Meanwhile, data limitations precluded inclusion of history of self-reported elevated blood glucose (Table S1). At baseline, ARIC study staff asked participants specifically about the use of glucose-lowering drugs and assessed fasting blood glucose of all participants, but did not inquire about gestational diabetes or history of other elevated blood glucose readings.

Although participants were asked to report parental history of diabetes at baseline, participants were not asked whether grandparents, siblings, or children had been diagnosed with diabetes. Thus, participants were assigned 5 points on the modified FINDRISC score if their biological mother or father ever had, or now has, diabetes (Table S1).

Data analysis

We used logistic regression to estimate odds ratios and 95% confidence intervals (OR, 95% CI) for the association between the modified FINDRISC and incident diabetes. To investigate the utility of the score to predict the outcome of interest, the modified FINDRISC was modeled as a continuous variable, and ROC curves were generated by plotting the sensitivity of the score versus the false-positive rate (1-specificity) for the overall study sample. Stratified analyses were then conducted according to race/gender groups to evaluate the consistency of the predictive properties of the modified FINDRISC for white and black men and women. We reported area under the curve (AUC) statistics for the study population, overall and by race/gender group.

To investigate the magnitude of association between each individual FINDRISC component and the outcome of incident diabetes (overall and by race/gender group), we also modeled the modified FINDRISC components separately as categorical variables. This analysis allowed us to understand the individual contribution of each of the components in the presence of the other factors. In an ad hoc analysis, we calculated AUCs and confidence intervals (CIs) for waist circumference and BMI alone, as categorical variables, due to the magnitude of the ORs for these FINDRISC components. Analyses were done using SAS version 9.2 (SAS institute, Inc. Cary NC.).

RESULTS

Baseline characteristics of ARIC study participants stratified by incident diabetes status at the end of 9 years follow-up are given in Table 1. The overall 9-year cumulative incidence of diabetes was 12.1% (1180/9754). By race/gender group, the cumulative incidence of diabetes was 20.3% for black women, 18.8% for black men, 12.4% for white men, and 8.6% for white women (data not shown; can be calculated from events and non-events in Table 3). Among individuals who developed diabetes, 49% were obese, over 70% had a high waist circumference, 40% reported use of anti-hypertensive medication at baseline, and 38% had a parental history of diabetes (Table 1). The mean modified FINDRISC was 10.5 (SD 3.3) among those who developed diabetes and 7.8 (SD 3.6) among those who did not.

The baseline characteristics of the study population by race/gender groups are given in Table 2. Forty-two percent of black women were obese and 70% had a high waist circumference, while only 27% of black men were obese and 28% had a high waist circumference (Table 2). Twenty-one percent of whites were obese, but gender differences in high waist circumference were observed: 63% of women versus 30% of men had a high waist circumference. The mean modified FINDRISC for black women was also higher than that of black men, white women, and white men at baseline. Antihypertensive medication use at baseline was common among all gender/by race subgroups (22%–44%), and was highest among black women.

Figure 1 represents the risk of incident diabetes for each race/gender subgroup by decile of the modified FINDRISC. The risk of diabetes generally increased in a dose-response manner across FINDRISC deciles for all race/gender groups. With the exception of white women, the distributions displayed an inflection point (increase) at the sixth decile, corresponding to

a modified FINDRISC of 9 (Figure 1). We also observed that within the same decile of the FINDRISC the magnitude of risk varied considerably across race/gender groups. Of note, the number of eligible ARIC participants was lower for blacks than for whites. Therefore, there were relatively small numbers of incident diabetes cases among blacks in each decile of risk which may have resulted in unstable estimates.

The ROC curve statistics for the continuous FINDRISC, overall and by race/gender groups, are given in Table 3. The area under the curve (AUC) for the risk of incident diabetes was highest for white women (0.77) and lowest for black men (0.70). We also modeled the individual FINDRISC components as categorical variables and observed multivariable ORs and 95% CIs for incident diabetes as shown in Table 4. Overall, the odds of diabetes were highest for obese study participants (4.00, 3.15–5.07), controlling for age, waist circumference, antihypertensive medication, and family history of diabetes.

Among black women (2.39, 1.00–5.75), black men (4.65, 1.96–11.0) and white men (3.13, 1.98–4.96), high waist circumference was the most dominant predictor of incident diabetes (Table 4). Meanwhile, obese BMI (3.42, 2.33–5.02) had the largest effect on risk for white women (Table 4). Family history of diabetes had a smaller magnitude of effect among black men compared to other race/gender groups (Table 4).

The ad hoc analyses of waist circumference and BMI yielded AUCs of 0.72 (0.71, 0.74) for waist circumference alone, and 0.73 (0.71, 0.74) for BMI alone in the overall sample (data not shown). Meanwhile, high waist circumference alone (5.08, 4.09–6.31) was associated with incident diabetes in the overall sample, and the magnitude of effect was highest among white women (7.67, 4.62–12.7) and lowest for black women (3.75, 1.88–7.50). Obese BMI alone (7.05, 5.85–8.48) was associated with incident diabetes in the overall sample, and its effect was highest for white women and lowest for black women (Table S2).

DISCUSSION

We used a modified FINDRISC algorithm to predict incident diabetes in the ARIC cohort among the subset of the original cohort without missing diabetes status at all follow-up visits. In this population, the risk of diabetes generally increased with an increase in the FINDRISC for all race/gender groups. At some deciles of the modified FINDRISC algorithm, the scores appear to represent different levels of absolute risk within each race/ gender group. This difference was most prominent for black women in the highest decile of risk, corresponding to a modified FINDRISC of 14 to17. Thus, we found that while the FINDRISC is predictive within race/gender groups, it does not appear from these data that there is a single threshold for absolute risk that can be applied to a diverse population. This is in keeping with previous findings in other populations.³⁰

Scores such as the FINDRISC may be especially useful for identifying people at-risk for diabetes outside of traditional medical care settings, since existing diabetes risk prediction algorithms that require fasting or 2-hour glucose measures also limit the contexts in which they can be applied. Prediction of diabetes with these non-clinical algorithms has been shown in most cases to improve only marginally beyond algorithms composed of fasting or

post-prandial glucose measures plus more well-established and clinically-available risk factors for diabetes. For the first time, we used a modified FINDRISC score for diabetes prediction among a diverse US-based population.

We analyzed the impact of each FINDRISC component in multivariable analyses, and found different magnitudes of effect for the components by race/gender group. For example, waist circumference was more predictive of incident diabetes than BMI among black men, but not among white women. In the context of using the FINDRISC algorithm as part of public health screening efforts to identify populations at greater risk for diabetes, our results from this subset of the original ARIC cohort indicate that either different threshold cut-points for different race and gender groups may be needed, or that for optimal utility, the algorithm may need to be calibrated to derive different metrics for the individual components for different race-, gender- or race-by-gender groups in order to derive a common score that represents a similar magnitude of risk for all groups.

Our results indicated that the modifiable measures of adiposity (BMI and waist circumference) were associated with higher odds of diabetes than other risk score components. Consistent with previous analyses in ARIC³⁰ and other population-based studies, we demonstrated that obesity and high waist circumference were critical for the prediction of diabetes but that age, use of anti-hypertensive medication, and family history of diabetes were not as strong of predictors for incident diabetes in this population. The AUCs for waist circumference (0.72) and BMI alone (0.73), were almost comparable to the modified FINDRISC (0.74) for the overall sample. Although this was true for our study sample, we do not have the data to determine whether this finding would be externally valid for other populations. It is possible that these observed relationships are due to the specific measures of adiposity and their correlations with other co-occurring factors not controlled for in our study sample. For example, the prevalence of obesity in our study population was higher compared to previous populations in which the FINDRISC was evaluated, including German²⁹ and US-representative²⁵ populations.

An unmodified FINDRISC was recently evaluated in a US population using NHANES ²⁵ data. Notably, the overall AUC (0.74 for ARIC versus 0.75 for NHANES) was similar, yet race- (0.74 for ARIC whites, 0.71 for ARIC blacks versus 0.76 for NHANES whites and blacks) and gender- (0.70 for men and 0.77 for women in ARIC versus 0.74 for men and 0.78 for women in NHANES) specific estimates differed slightly, using undiagnosed diabetes as the endpoint of comparison in NHANES. There were some key differences between these investigations: our analysis was longitudinal and assessed incident diabetes, while the NHANES analysis was cross-sectional and identified participants with undiagnosed diabetes and pre-diabetes.²⁵ Also, the NHANES analysis used the original FINDRISC to evaluate diabetes while we used a modified FINDRISC in the current study.

Our study included a large, multicenter, biracial population for analysis. To our knowledge, the current investigation using data from the ARIC study represents the first time the FINDRISC has been used for cross-temporal prediction of incident diabetes among black and white individuals living in the US. A possible limitation of the FINDRISC in our study population is that the metric does not differentiate between obesity and morbid obesity (BMI

>40 kg/m²), even though there may be a difference in the incidence of diabetes for obese versus morbidly obese participants.

Although the original FINDRISC score includes simple measures of physical activity (30 minutes a day on most days) and dietary patterns (fruit and vegetable consumption), developers of the algorithm have clarified that neither item added much to the predictive power of the statistical model, but were included in the risk score for public health purposes to emphasize the importance of physical activity and diet in the prevention of diabetes.³¹ Thus, not utilizing these two measures in the ARIC data likely did not affect the diagnostic properties of the modified FINDRISC in this population. However, it should be acknowledged that we also were not able to account for prior history of elevated glucose/ gestational diabetes, nor extended family history of diabetes. Limitations of our data source may have precluded the characterization of conditions such as pre-diabetes, which may contribute to the predictive power of the FINDRISC in certain race/gender groups.

A limitation of using these ARIC data to re-calibrate the FINDRISC score, as previously described, is the differential loss to follow-up of persons of minority race/gender groups. Of note, these same groups have a smaller representation to begin with, have competing risks which may have precluded their eligibility for these analyses, and may be more likely to be missing diabetes status during follow-up.

Regardless of these limitations, the modified FINDRISC may be useful for the early screening of incident diabetes in biracial populations. Importantly, the score could be used as a tool for informing the public about risk factors associated with diabetes in order to help them to make healthy lifestyle choices to address obesity. Such an assessment of diabetes risk may allow for early intervention to delay or prevent diabetes. Finally, caution should be employed when interpreting this type of diabetes prediction model in biracial populations as we found differential prediction by race.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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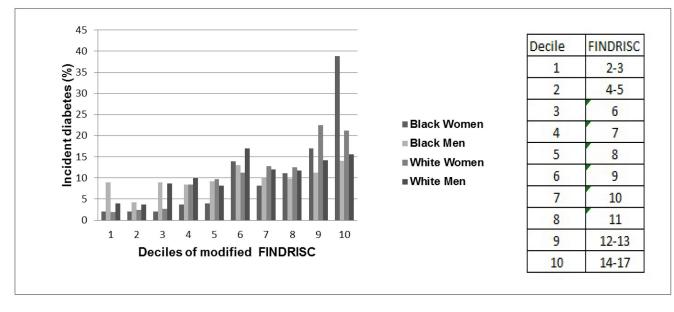
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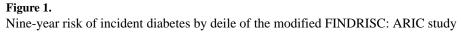


Table 1

Baseline characteristics (n, %) of participants with and without incident diabetes over 9 years' follow-up: ARIC study.

	Diabetes N=1,180	No Diabetes N=8,574
Age		
45–54 years	646 (54.8)	4,705 (54.9)
55–64 years	534 (45.2)	3,869 (45.1)
Gender		
Female	676 (51.1)	4,829 (56.3)
Male	647 (48.9)	3,745 (43.7)
Race		
Black	351 (29.7)	1,425 (16.6)
White	829 (70.3)	7,149 (83.4)
Study community		
Forsyth	279 (23.6)	2,247 (26.2)
Jackson	301 (25.5)	1,269 (14.8)
Minnesota	257 (21.8)	2,659 (31.0)
Washington	343 (29.1)	2,399 (28.0)
Body mass index (BMI)		
Normal <25 kg/m ²	154 (13.1)	3,359 (39.1)
Overweight 25- 30 kg/m ²	451 (38.2)	3,504 (40.9)
Obese >30 kg/m ²	575 (48.7)	1,711 (20.0)
Waist circumference		
Low (Women <80cm; Men <94cm)	96 (8.1)	2,177 (25.4)
Medium (Women 80-88cm; Men 94-102cm)	231 (19.6)	2,696 (31.4)
High (Women >88cm; Men >102cm)	853 (72.3)	3,701 (43.2)
Hypertensive medication		
Yes	477 (40.4)	2,037 (23.8)
No	703 (59.6)	6,537 (76.2)
Parental history of diabetes		
Yes	446 (37.8)	1,989 (23.2)
No	734 (62.2)	6,585 (76.8)
Modified FINDRISC score (mean, SD)	10.5 (3.3)	7.8 (3.5)

Table 2

Distributions (n, %) of the modified FINDRISC score and its individual components: ARIC study

	Black Women (n=1,149)	Black Men (n=627)	White Women (n=4,282)	White Men (n=3,696)
Age (years)				
45–54	745 (64.8)	393 (62.7)	2,375 (55.5)	1,838 (49.7)
55–64	404 (35.2)	234 (37.3)	1,907 (44.5)	1,858 (50.3)
Body mass index, kg/m ²				
<25	237 (20.6)	185 (29.5)	2,070 (48.3)	1,021 (27.6)
25- 30	431 (37.5)	272 (43.4)	1,336 (31.2)	1,916 (51.8)
>30	481 (41.9)	170 (27.1)	876 (20.5)	759 (20.6)
Waist circumference, cm				
Low (Women/<80; Men/<94)	116 (10.1)	274 (43.7)	819 (19.1)	1,064 (28.8)
Medium (Women/80-88; Men/94-102)	225 (19.6)	177 (28.2)	1,101 (25.7)	1,424 (38.5)
High (Women/>88; Men/>102)	808 (70.3)	176 (28.1)	2,362 (55.2)	1,208 (32.7)
Antihypertensive medications				
Yes	506 (44.0)	191 (30.5)	1,010 (23.6)	807 (21.8)
No	643 (56.0)	436 (69.5)	3,272 (76.4)	2,889 (78.2)
Family history of diabetes				
Yes	379 (33.0)	178 (28.4)	1,044 (24.4)	834 (22.6)
No	770 (67.0)	449 (71.6)	3,238 (75.6)	2,862 (77.4)
Modified FINDRISC score				
Mean (SD)	9.9 (3.6)	7.6 (3.9)	8.0 (3.6)	7.7 (3.5)

Table 3

Area under the curve (AUC) statistics for the modified FINDRISC to predict incident diabetes among ARIC participants during 9-year follow-up

	AUC (95% CI)	Event	Non-event
Overall	0.74 (0.72, 0.75)	1,180	8,574
Black Women	0.71 (0.68, 0.75)	233	916
Black Men	0.70 (0.65, 0.75)	118	509
White Women	0.77 (0.74, 0.79)	369	3,913
White Men	0.71 (0.68, 0.73)	460	3,236

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Table 4

The association (OR * 95% CI) between the individual components of the modified FINDRISC and incident diabetes, overall and by race/gender group: ARIC study.

	Overall	African American Women	African American Men	Caucasian Women	Caucasian Men
Age (years) 45-54 55-64	1 (referent) 0.99 (0.87, 1.12)	1 (referent) 0.92 (0.67, 1.25)	1 (referent) 0.92 (0.60, 1.41)	1 (referent) 1.10 (0.88, 1.38)	1 (referent) 0.94 (0.77, 1.16)
Body mass index (kg/m ²) Normal <25 Overweight 25- 30 Obese >30	1 (referent) 2.04 (1.65, 2.53) 4.00 (3.15, 5.07)	1 (referent) 0.92 (0.48, 1.78) 1.62 (0.83, 3.14)	l (referent) 0.95 (0.45, 2.02) 1.13 (0.44, 2.96)	1 (referent) 1.97 (1.37, 2.85) 3.42 (2.33, 5.02)	1 (referent) 1.21 (0.82, 1.78) 2.50 (1.56, 4.00)
Waist circumference (cm) Low (Men <94; Women <80) Medium (Men 94- 102; Women 80- 88) High (Men >102; Women >88)	1 (referent) 1.34 (1.03, 1.74) 1.92 (1.47, 2.50)	l(referent) 0.69 (0.27, 1.74) 2.39 (1.00, 5.75)	1 (referent) 2.27 (1.11, 4.64) 4.65 (1.96, 11.0)	1 (referent) 1.40 (0.77, 2.57) 2.82 (1.56, 5.10)	1 (referent) 2.09 (1.40, 3.14) 3.13 (1.98, 4.96)
Antihypertensive medication No Yes	1 (referent) 1.61 (1.41, 1.84)	1 (referent) 1.42 (1.06, 1.92)	l (referent) 1.24 (0.81, 1.90)	1 (referent) 2.29 (1.82, 2.87)	1 (referent) 1.32 (1.05, 1.66)
Family history of Diabetes No Yes	1 (referent) 1.85 (1.62, 2.11)	1 (referent) 2.10 (1.56, 2.82)	1 (referent) 1.27 (0.82, 1.97)	1 (referent) 1.89 (1.50, 2.38)	1 (referent) 1.89 (1.52, 2.34)

Diabetes Obes Metab. Author manuscript; available in PMC 2018 September 01.

 * Odds ratios are adjusted for all other variables in the table.