

Recto running head: Undiagnosed diabetes algorithms and nomogram

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## Original Article

### Simple non-laboratory- and laboratory-based risk assessment algorithms and nomogram for detecting undiagnosed diabetes mellitus

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## Abstract

**Background:** The aim of the present study was to develop a simple nomogram that can be used to predict the risk of diabetes mellitus (DM) in the asymptomatic non-diabetic subjects based on non-laboratory- and laboratory-based risk algorithms.

**Methods:** Anthropometric data, plasma fasting glucose, full lipid profile, exercise habits, and family history of DM were collected from Chinese non-diabetic subjects aged 18–70 years. Logistic regression analysis was performed on a random sample of 2518 subjects to construct non-laboratory- and laboratory-based risk assessment algorithms for detection of undiagnosed DM; both algorithms were validated on data of the remaining sample (n = 839). The Hosmer–Lemeshow test and area under the receiver operating characteristic (ROC) curve (AUC) were used to assess the calibration and discrimination of the DM risk algorithms.

**Results:** Of 3357 subjects recruited, 271 (8.1%) had undiagnosed DM defined by fasting glucose  $\geq 7.0$  mmol/L or 2-h post-load plasma glucose  $\geq 11.1$  mmol/L after an oral glucose tolerance test. The non-laboratory-based risk algorithm, with scores ranging from 0 to 33, included age, body mass index, family history of DM, regular exercise, and uncontrolled blood pressure; the laboratory-based risk algorithm, with scores ranging from 0 to 37, added triglyceride level to the risk factors. Both algorithms demonstrated acceptable calibration (Hosmer–Lemeshow test:  $P = 0.229$  and  $P = 0.483$ ) and discrimination (AUC 0.709 and 0.711) for detection of undiagnosed DM.

**Conclusion:** A simple-to-use nomogram for detecting undiagnosed DM has been developed using validated non-laboratory-based and laboratory-based risk algorithms.

- Significant findings of the study: Validated non-laboratory- and laboratory-based risk assessment algorithms were developed for the prediction of undiagnosed DM in a Chinese population. The optimal cut-off point on the ROC curve was 18 for the detection of undiagnosed DM in both algorithms.
- What this study adds: The simple and user-friendly nomogram was constructed to enable clinicians to estimate an individual's risk of DM and thus to promote targeted screening for DM among high-risk individuals in the primary care and community settings.

**Key words:** calibration, discrimination, nomogram, risk algorithm, undiagnosed diabetes, validation.

## <A>Introduction

Diabetes mellitus (DM) is highly prevalent and has become one of the major disease burdens worldwide.<sup>1</sup> In particular, undiagnosed DM is associated with a higher risk of diabetes-related complications<sup>2</sup> and mortality<sup>3</sup> compared with normal glucose tolerance because of the lack of awareness of high blood glucose levels and delayed disease management. Early detection of DM through periodic screening has been recommended<sup>4</sup> for high-risk individuals using 75-g oral glucose tolerance tests (OGTT), fasting plasma glucose (FPG) tests or HbA1c levels. In a recent cost-effectiveness analysis, screening for DM and prediabetes was found to be cost-saving among patients at risk compared with no screening.<sup>5</sup> Nevertheless, a targeted screening approach based on risk assessment and stratification of high-risk subjects should be undertaken to optimize resource allocation and utilization.<sup>6</sup>

In order to stratify high-risk subjects for DM screening, risk algorithms for the detection of undiagnosed DM have been developed and validated in different populations and healthcare settings.<sup>6-9</sup> However, these risk score algorithms are population specific; the combination of risk factors in each algorithm varies across countries, ethnicities, and levels of income based on the population in which these algorithms were developed. For the Chinese population,<sup>10-13</sup> most of the risk algorithms<sup>11-13</sup> were derived from results of laboratory tests, in addition to sociodemographic and anthropometric data. Interestingly, the majority of these Chinese DM risk algorithms were developed and validated in low income settings.<sup>10-12</sup> Little is known about their performance, calibration, and discrimination in the detection of undiagnosed DM among Chinese populations in non-low income settings. Furthermore, no nomograms for the identification of undiagnosed DM have been developed, which would be an ideal, simple-to-use graphical tool to facilitate DM risk assessment and stratification in clinical practice and the community.

y setting.

The aims of this study were to present and validate a non-laboratory- and laboratory-based risk assessment algorithm for detecting undiagnosed DM among the Chinese population in a non-low income setting. In addition, a simple and user-friendly nomogram was constructed to enable clinicians to estimate individual DM risk based on non-laboratory and laboratory risk assessment algorithms. Both tools could inform clinical decision making and enable identification of individuals at high risk for DM to undergo further screening tests, thus promoting the early detection of DM in the clinical and community settings. In addition, the study provides information on modifiable risk factors to be targeted by health intervention programs for the reduction of individual DM risk.

## **<A>Methods**

### *<B>Subjects for the development of risk assessment algorithms*

This study was part of a Hong Kong professional driver community project to promote health awareness and literacy regarding DM. Details regarding subject recruitment, eligibility criteria, and interviews have been reported previously.<sup>14–16</sup>

Subjects were excluded if they self-reported to have clinician-diagnosed DM. For each eligible subject, we retrieved DM risk factor data, including sociodemographic characteristics, lifestyle, past medical history and family history, which were surveyed through structured interview questionnaires. Sociodemographic data included the age, gender, and marital status of the subjects. For lifestyle factors, we considered self-reported drinking status, smoking status, and exercise frequency. For past medical history, gestational DM was defined by self-reported high blood glucose levels during pregnancy in women without a known diagnosis of DM. A self-reported history of hypertension was recorded. A positive family history of DM was defined as either first-degree (parents or siblings) or second-degree (grandparents) relatives having DM.

In addition, anthropometric and laboratory assessments were performed on each subject at baseline. For anthropometric assessment, we measured body mass index (BMI), systolic blood pressure (SBP), diastolic blood pressure (DBP), and waist circumference. Blood samples were taken for the measurement of FPG and 2-h post-load plasma glucose in the 75-g OGTT to determine glycemic status, as well as to determine the full lipid profile, including triglycerides (TG), total cholesterol (TC), high-density lipoprotein cholesterol (HDL-C), and low-density lipoprotein cholesterol (LDL-C), to assess DM risk. Plasma glucose, TG, TC, and HDL-C were measured using the Abbott Architect c16000 chemistry analyzer, whereas LDL-C was derived from the Friedewald formula.

Undiagnosed DM was defined as subjects with FPG  $\geq 7.0$  mmol/L or 2-h post-load plasma glucose  $\geq 11.1$  mmol/L in the 75-g OGTT.

#### *<B>Development and validation of risk assessment algorithms*

In all, 3357 subjects completed the questionnaire survey and underwent anthropometric and laboratory assessment during the study period. Simple random sampling was performed to select 2518 subjects from the total number of subjects (75% of the total sample) as the development sample. The remaining 25% formed the validation sample.

Data from the development sample (n = 2518) on the following DM risk factors were used to develop the risk assessment model: (i) sociodemographic and lifestyle risk factors, including age (<45,  $\geq 45$ –<50,  $\geq 50$ –<55,  $\geq 55$ –<60, and  $\geq 60$  years), gender, smoking status (active smoker), and exercise frequency; (ii) non-laboratory clinical risk factors, including family history of DM, history of hypertension, obesity (BMI  $\geq 25$  kg/m<sup>2</sup>), central obesity (waist circumference  $\geq 80$  cm for women and  $\geq 90$  cm for men), and uncontrolled blood pressure (SBP  $\geq 140$  mmHg or DBP  $\geq 90$  mmHg); and (iii) laboratory risk factors, including TG, TC, HDL-C, and LDL-C. Sociodemographic and non-laboratory clinical risk factors were used for the development of the non-laboratory-based risk assessment algorithm. Laboratory risk factors were added to

sociodemographic and non-laboratory clinical risk factors for the development of the laboratory-based risk assessment algorithm.

Based on undiagnosed DM as the outcome, significant risk factors with  $P < 0.05$  in a stepwise binary logistic regression model were retained in the final risk assessment model. Each risk factor was assigned a weighting in the risk score using respective  $\beta$ -coefficients multiplied by 10 and rounded to the nearest integer. The risk score for each subject would be the sum of risk score contributed by each risk factor identified by the final risk assessment model.

Non-laboratory- and laboratory-based risk assessment algorithms were validated externally using data for the remaining 839 subjects in the validation sample; these subjects had not been used for algorithm development. The accuracy, calibration, and discrimination of the risk algorithms to detect undiagnosed DM were compared against six previously published DM risk assessment algorithms,<sup>10,12,13, 17–19</sup> of which three were developed in Chinese populations (New Chinese Diabetes Risk Score,<sup>10</sup> Qingdao Diabetes Risk Score,<sup>12</sup> and Southern Chinese Risk Score<sup>13</sup>).

#### **<B>***Design of a nomogram*

Both risk assessment algorithms were then converted into a nomogram (Fig. 1). With regard to the design of the nomogram, the score for each characteristic for each subject was mapped directly on the nomogram using the rule located at the top of the nomogram (Fig. 1). The scores obtained for each characteristic are summed to compute the total DM risk score based on the risk assessment algorithm. Total DM risk scores are mapped in the lower part of the nomogram to predict the prevalence of undiagnosed DM.

#### **<B>***Data analysis*

Descriptive statistics of baseline characteristics between development and validation samples were compared using independent  $t$ -tests for continuous variables and Chi-squared tests for ca

tegorical variables.

Risk algorithms were developed and modeled using a stepwise binary logistic regression model. The goodness-of-fit of the two models was assessed by Akaike information criterion (AIC) and Bayesian information criterion (BIC) statistics, with the lower values indicating better fit. For each risk algorithm, Youden's index was used to determine the optimal cut-off value for detecting undiagnosed DM.<sup>20</sup> Accuracy in terms of sensitivity and specificity of the use of risk algorithms at optimal cut-off values for detecting undiagnosed DM were compared with diagnosis confirmed by FPG or OGTT as the diagnostic standard. The Hosmer–Lemeshow  $\chi^2$  statistic and area under the receiver operating characteristic (ROC) curve (AUC) were used to assess the calibration and discrimination of these risk algorithms.

All analyses were conducted using SPSS version 20.0 for Windows (IBM Inc., Chicago, IL, USA) and STATA version 13 (STATA Corporation, College Station, TX, USA). Two-sided  $P < 0.05$  was considered significant.

## **<A>Results**

Descriptive characteristics of the 3357 study subjects overall and by analysis samples are presented in Table 1. Of 3357 subjects without a prior history of DM, 271 (8.1%) had undiagnosed DM. The prevalence of undiagnosed DM between the development and validation samples did not differ significantly (8.3% vs 7.3%;  $P = 0.325$ ). The mean ( $\pm$  SD) age of subjects was  $50.9 \pm 7.6$  years. Most subjects were male (92.7%) and non-smokers (80.5%). In terms of non-laboratory-based clinical characteristics, 31.4% of subjects had a family history of DM, 51.2% had general obesity, 48.1% had central obesity, and 32.2% had uncontrolled blood pressure with either SBP  $>140$  mmHg or DBP  $>90$  mmHg; 12.8% of subjects had history of hypertension and the subjects in the validation sample were more likely to have hypertension than those in the development sample. There were no significant differences in laboratory-based clinical characteristics between the development and validation samples.

For model development based on stepwise binary logistic regression (non-laboratory- vs laboratory-based), there was a significant association between undiagnosed DM and increasing age for each 5-year stratum from 45 to 60 years ( $\beta$ -coefficient 0.49–1.10 vs 0.19–1.12), general obesity ( $\beta$ -coefficient 0.45 vs 0.35), a positive family history of DM in a first- or second-degree relative ( $\beta$ -coefficient 0.72 vs 0.70), exercising regularly ( $\beta$ -coefficient 0.60 vs 0.57), and uncontrolled blood pressure ( $\beta$ -coefficient 0.54 vs 0.052). Abnormal TG was significantly associated with undiagnosed DM in the regression analysis for the laboratory-based risk assessment algorithm. Significant risk factors were used to assign weighted scores, and the total risk score for each subject was the sum of all risk scores allocated to each individual risk factor. The scores for non-laboratory- and laboratory-based risk assessment algorithms ranged from 0 to 33 and from 0 to 37, respectively. Youden's index suggesting the optimal cut-off value for undiagnosed DM was 18 for both risk assessment algorithms. At the optimal cut-off value  $\geq 18$ , the sensitivity and specificity were 57.9% and 68.9%, respectively, for the non-laboratory-based algorithm and 66.2% and 60.2%, respectively, for the laboratory-based algorithm.

For the external validation of models, the AUC for the non-laboratory- and laboratory-based algorithms was 0.709 and 0.711, respectively, supporting model discrimination. The Hosmer–Lemeshow test with  $P > 0.05$  (0.229 and 0.483, respectively) indicated adequate calibration of the non-laboratory- and laboratory-based algorithms. Using the validation sample, the sensitivity and specificity at the optimal cut-off value  $\geq 18$  for the non-laboratory-based risk algorithm were 63.9% and 67.7%, respectively, whereas those of the laboratory-based risk algorithm were 72.1% and 57.8%, respectively.

Compared with existing risk algorithms, the risk algorithms developed in the present study exhibited adequate accuracy, discrimination, and calibration. Figure 1 shows the nomogram that graphically calculated the non-laboratory- and laboratory-based risk scores. The lower part of the nomogram shows the predicted probability of the individual having undiagnosed DM in both routine clinical practice and the community setting. From the nomogram, the predic

ted prevalence of undiagnosed DM increases gradually from 1% at a total risk score of 0 to 34% at a total risk score of 33.

## **<A>Discussion**

The present study developed and validated simple non-laboratory- and laboratory-based risk assessment algorithms for predicting undiagnosed DM in the general Chinese population. Based on the risk assessment algorithms, we developed a simple-to-use nomogram (Fig. 1) for primary care clinicians to facilitate risk sharing in such a way that high-risk subjects are identified to promote the uptake of DM screening. In addition to risk information sharing, this risk assessment facilitates the health service provider prioritizing DM prevention strategies and launching DM screening in the primary care setting when resources for glycemic testing are limited. Moreover, risk stratification identified high-risk subjects, who were then encouraged to undergo DM screening using the most appropriate screening test. Screened subjects were empowered to increase their awareness of DM and to make lifestyle modifications.

Risk scores with cut-off values  $\geq 18$  had the best combination of sensitivity and specificity for detecting undiagnosed DM. Therefore, subjects with risk scores  $\geq 18$  were considered to be high-risk subjects recommended to undergo DM screening. Visualization of the nomogram has important implications for clinicians and subjects. For example, the predicted probability of undiagnosed DM is approximately 15% if subjects score 23 on the non-laboratory-based risk assessment algorithm.

The National Prevalence Health Survey of 46 239 adults in 14 provinces in China found that the prevalence of undiagnosed DM was 6.3%,<sup>21</sup> and a survey conducted in a nationally representative sample of 98 658 Chinese adults reported a prevalence of 8.1% in 2010.<sup>22</sup> The prevalence of undiagnosed DM found in the development (8.3%) and validation (7.3%) samples in the present study is comparable to that in the Chinese population. Thus, the prevalence of undiagnosed DM in the present study was comparable to the Hong Kong prevalence of 9.51%,

as estimated by the International Diabetes Federation.<sup>23</sup>

The non-laboratory-based risk assessment algorithm was constructed on the basis of five widely recognized risk factors: age, BMI, family history of DM, exercise frequency, and blood pressure. This is in line with recent systematic reviews<sup>7,8</sup> of newly developed risk assessment algorithms reporting that the frequently included risk factors are age and BMI, representing measures of body mass. Age group was a main contributor to the risk score for the detection of undiagnosed DM, because we observed a trend for increasing risk score with increasing age. Interestingly, there were two modifiable risk factors recognized in our algorithms: BMI and exercise frequency. This implies that health intervention programs, such as body weight monitoring and control and lifestyle interventions, are useful in reducing DM risk in asymptomatic subjects in the Chinese population.

Although the risk factors identified in our algorithms are mostly found in existing algorithms summarized by systematic reviews,<sup>7,8</sup> there are no existing risk algorithms using the same pool of risk factors. Risk factors may be presented in other forms and using alternative definitions. It should be highlighted that there is no universal consensus regarding the combination of risk factors to be used for the detection of undiagnosed diabetes. Notably, the risk algorithms developed in one country may not be transferable to other countries, reflected by the poor ability to calibrate and discriminate the external dataset. For example, two risk algorithms<sup>12,13</sup> developed using the Chinese population did not have satisfactory performance, with an AUC of  $\geq 0.7$  in our dataset upon external validation; this could be due, in part, to differences in income levels. However, recent studies<sup>24,25</sup> have not found any clear indication that the addition of ethnicity improves the performance of risk algorithms for predicting diabetes. Therefore, further research is required to improve the performance through the additional effects of alternative key information, such as spousal history of diabetes.<sup>26</sup>

A high degree of model discrimination does not necessarily imply identification of true positive diabetes cases. Upon external validation of existing algorithms developed using Cau

asian populations, the AUC of the algorithm of Bang et al. was  $<0.7$ .<sup>17</sup> Even though the discrimination of another two algorithms<sup>18,19</sup> was greater than that of our algorithms, those algorithms did not achieve a good trade-off between sensitivity and specificity at predefined optimal cut-off values. For both algorithms,<sup>18,19</sup> the specificities were unacceptably low although the sensitivities were at least 90%. Hence, a high degree of discrimination does not necessarily lead to reasonable performance at recommended cut-off values. In such cases, when an algorithm has a high degree of discrimination upon external validation, new recommended cut-off values may help balance sensitivity and specificity.

The laboratory-based risk assessment algorithm increased the number of risk factors by including factors such as TG; however, this algorithm may not be able to stratify risks of asymptomatic subjects without prior laboratory testing. Nevertheless, such an algorithm is likely to be used when identifying high-risk subjects in the routine clinical setting, where blood samples are routinely collected for lipid profile assessment.

### **<B>***Limitations*

Several limitations of present study should be noted. First, our risk algorithms were developed and validated using Chinese data, so they may not be generalizable to non-Chinese populations. However, only two previous studies<sup>27,28</sup> included ethnicity in their algorithms, indicating that ethnicity is not a common predictive factor for the prediction of diabetes. Second, there was a high proportion (92.7%) of male professional drivers in our development and validation samples.

### **<B>***Conclusions*

In summary, validated risk assessment algorithms were developed for the prediction of undiagnosed DM in a Chinese population. A corresponding simple-to-use nomogram was constructed to facilitate risk information sharing and to promote targeted screening for DM among high-

risk individuals in the primary care and community settings. Future studies on the evaluation of our risk algorithms and nomogram to examine the feasibility and validity of their use in targeted community-based screening for DM are warranted.

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### **<A>Disclosure**

None declared.

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**Figure 1** Nomogram to predict the probability of undiagnosed diabetes based on non-laboratory- and laboratory-based risk algorithms. The patient’s score for each parameter is plotted on the appropriate scale and vertical lines are drawn to the line of points to obtain the corresponding scores. All scores are summed to obtain a total points score. The total points score is plotted on the total points line and a vertical line is drawn down to the bottom line. The corresponding value shows the predicted probability of undiagnosed diabetes.

**Table 1** Sociodemographic and clinical characteristics of subjects without a known history of diabetes in the model development and validation samples

	Total (n = 3357)	Model development sample (n = 2518)	Model validation sample (n = 839)	P-value
Sociodemographic characteristics				
Age (years)				0.113
<45	637 (19.0%)	501 (19.9%)	136 (16.2%)	
≥45 and <50	767 (22.8%)	571 (22.7%)	196 (23.4%)	
≥50 and <55	789 (23.5%)	595 (23.6%)	194 (23.1%)	
≥55 and <60	747 (22.3%)	541 (21.5%)	206 (24.6%)	
≥60	417 (12.4%)	310 (12.3%)	107 (12.8%)	
Gender				0.816
Female	246 (7.3%)	183 (7.3%)	63 (7.5%)	

Male	3111 (92.7%)	2335 (92.7%)	776 (92.5%)	
Exercise regularly	1480 (44.1%)	1089 (43.3%)	391 (46.6%)	0.092
Smoking	653 (19.5%)	485 (19.3%)	168 (20.0%)	0.629
Clinical characteristics				
Undiagnosed diabetes	271 (8.1%)	210 (8.3%)	61 (7.3%)	0.325
Family history of diabetes	1053 (31.4%)	778 (30.9%)	275 (32.8%)	0.310
Gestational diabetes <sup>A</sup>	19 (7.7%)	17 (9.3%)	2 (3.2%)	0.117
Hypertension	429 (12.8%)	305 (12.1%)	124 (14.8%)	0.045*
WC (cm)	89.1 ± 8.2	89.0 ± 8.3	89.5 ± 8.2	0.105
Central obesity (WC ≥ 90 cm men; ≥ 80 cm women)	1616 (48.1%)	1196 (47.5%)	420 (50.1%)	0.198
BMI (kg/m <sup>2</sup> )	25.4 ± 3.4	25.3 ± 3.4	25.6 ± 3.4	0.094
General obesity (BMI ≥ 25 kg/m <sup>2</sup> )	1719 (51.2%)	1270 (50.4%)	449 (53.5%)	0.122
SBP (mmHg)	132.4 ± 15.9	132.2 ± 15.9	133.0 ± 16.1	0.242
DBP (mmHg)	80.3 ± 10.7	80.1 ± 10.7	80.8 ± 10.7	0.113
Suboptimal BP (SBP >140 or DBP >90 mmHg)	1082 (32.2%)	810 (32.2%)	272 (32.4%)	0.893
FPG (mmol/L)	5.4 ± 1.4	5.4 ± 1.5	5.3 ± 1.2	0.160
TG (mmol/L)	1.7 ± 1.3	1.7 ± 1.3	1.7 ± 1.1	0.997
TC (mmol/L)	5.4 ± 1.0	5.4 ± 1.0	5.4 ± 0.9	0.470
HDL-C (mmol/L)	1.2 ± 0.3	1.2 ± 0.3	1.2 ± 0.3	0.652
LDL-C (mmol/L)	3.5 ± 0.9	3.5 ± 0.9	3.5 ± 0.8	0.477

Data are given as the mean ± SD or as the number of subjects with percentages in parentheses. Asterisks indicate a significant difference ( $P < 0.05$ ) between the development and validation samples (independent  $t$ -test or Chi-squared test, as appropriate).

<sup>A</sup>Only for women (n = 246).

BMI, body mass index; BP, blood pressure; SBP, systolic BP; DBP, diastolic BP; TC, total cholesterol; TG, triglycerides; FPG, fasting plasma glucose; WC, waist circumference.

**Table 2** Risk factors and their respective risk scores for non-laboratory- and laboratory-based risk assessment algorithms based on the development sample

Factors	Non-laboratory-based algorithm				Laboratory-based algorithm					
	$\beta$ -Coefficient	OR	95% (CI)	$P$ -value	Score	$\beta$ -Coefficient	OR	95% (CI)	$P$ -value	Score
Age (years)										
<45	–	1	–	–	0	–	1	–	–	0
≥ 45 and <50	0.49	1.63	(0.98, 2.73)	0.061	5	0.49	1.63	(0.98, 2.73)	0.062	5
≥ 50 and <55	0.58	1.79	(1.08, 2.97)	0.024*	6	0.59	1.80	(1.09, 2.99)	0.022*	6

≥55 and <60	0.83	2.30 (1.39, 3.80)	0.001*	8	0.84	2.31 (1.39, 3.83)	0.001*	8
≥60	1.10	3.02 (1.74, 5.23)	<0.001*	11	1.12	3.06 (1.76, 5.30)	<0.001*	11
BMI (kg/m <sup>2</sup> )								
<25	–	1	–	0	–	1	–	0
≥25	0.45	1.56 (1.16, 2.11)	0.003*	4	0.35	1.43 (1.05, 1.94)	0.024*	4
Family history of diabetes								
No	–	1	–	0	–	1	–	0
Yes	0.72	2.05 (1.53, 2.75)	<0.001*	7	0.70	2.02 (1.51, 2.71)	<0.001*	7
Exercise regularly								
Yes	–	1	–	0	–	1	–	0
No	0.60	1.82 (1.34, 2.48)	<0.001*	6	0.57	1.76 (1.29, 2.41)	<0.001*	6
Suboptimal blood pressure								
No	–	1	–	0	–	1	–	0
Yes	0.54	1.71 (1.28, 2.30)	<0.001*	5	0.52	1.69 (1.25, 2.27)	0.001*	5
Triglyceride (mmol/L)								
<1.7		Not considered			–	1	–	0
≥1.7					0.43	1.53 (1.14, 2.07)	0.005*	4
Goodness-of-fit								
AIC		1, 381				1, 375		
BIC		1, 433				1, 433		

AIC, Akaike information criterion; BIC, Bayesian information criterion; BMI, body mass index; CI, confidence interval; OR, odds ratio.

**Table 3** Accuracy, discrimination and calibration of the risk algorithms developed in the present study and existing risk algorithms using the development (internal validation) and validation (external validation) samples

Risk algorithm	Optimal cut-off value (range)	AUC (95% CI)	Sensitivity (%) at cut-off value (95% CI)	Specificity (%) at cutoff value (95% CI)	Hosmer–Lemeshow test
Present study					
Internal validation (n = 2518)					
Non-laboratory-based algorithm	18 (0–33)	0.686 (0.650, 0.722)	57.9 (51.8, 63.9)	68.9 (67.3, 70.6)	0.159
Laboratory-based algorithm	18 (0–37)	0.696 (0.661, 0.731)	66.2 (59.4, 72.6)	60.2 (58.2, 62.2)	0.053
External validation (n = 839)					
Non-laboratory-based algorithm	18 (0–33)	0.709 (0.646, 0.773)	63.9 (50.6, 75.8)	67.7 (64.3, 71.0)	0.229
Laboratory-based algorithm	18 (0–37)	0.711 (0.648, 0.774)	72.1 (59.2, 82.9)	57.8 (54.3, 61.3)	0.483
Existing algorithms					
New Chinese Diabetes risk score [10]	25 (0–51)	0.708 (0.644, 0.772)	98.4 (91.2, 100.0)	15.0 (12.6, 17.7)	0.730
Qingdao Diabetes risk Score [12]	Men 17 (3–32) Women 14 (3–32)	0.672 (0.602, 0.742)	75.4 (62.7, 85.5)	53.1 (49.5, 56.6)	0.383
Southern Chinese Risk score [13]	16 (0–30)	0.664 (0.595, 0.734)	49.2 (36.1, 62.3)	75.2 (72.0, 78.2)	0.208
Spanish Diabetes Risk score [18]	100 (unspecified)	0.728 (0.669, 0.788)	100.0 (94.1, 100.0)	0.1 (0.0, 0.7)	0.935
Patient Self-Assessment score [17]	5 (-1–9)	0.675 (0.607, 0.743)	37.7 (25.6, 51.0)	82.5 (79.7, 85.1)	0.986
TOPICS Diabetes Screening score [19]	8 (0–16)	0.723 (0.663, 0.784)	91.8 (81.9, 97.3)	33.2 (29.9, 36.6)	0.320

NA, not applicable; AUC, area under the receiver operating characteristic curve; TOPICS, Toranomom Hospital Health Management Center Study; CI, confidence interval.