BMJ Open Incidence and risk factors of type 2 diabetes mellitus in an overweight and obese population: a long-term retrospective cohort study from a Gulf state

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ABSTRACT

Objectives A high body mass index (BMI) is associated with risk of type 2 diabetes mellitus (DM). The United Arab Emirates (UAE) is experiencing a marked increase in obesity. Nonetheless, no data are available regarding the incidence of type 2 DM in the high-risk adult UAE population. Therefore, this study aimed to evaluate the incidence rate and risk of developing type 2 DM among individuals with above-normal BMI in the UAE.

Design A retrospective cohort study.

Setting Outpatient clinics at a tertiary care centre in Al Ain, UAE.

Participants Three hundred and sixty-two overweight or obese adult UAE nationals who visited outpatient clinics between April 2008 and December 2008.

Primary outcome measure Patients with type 2 DM were identified based on diagnosis established by a physician or through glycated haemoglobin (HbA1c) levels \geq 6.5% during the follow-up period (until April 2018). **Results** The overall incidence rate of type 2 DM during the median follow-up time of 8.7 years was 16.3 (95% Cl 12.1 to 21.4) cases per 1000 person-years. Incidence rates in men and women were 17.7 (95% Cl 11.6 to 25.9)

and 15.0 (95% Cl 9.8 to 22.2) cases per 1000 personyears, respectively. Multivariable Cox proportional hazard analysis determined older age and obesity in women and pre-diabetes in men to be independent risk factors for developing type 2 DM.

Conclusions The incidence rate of type 2 DM in overweight and obese UAE nationals is high. In addition to screening, current strategies should strongly emphasise lifestyle modifications to decrease HbA1c and BMI levels in this high-risk population.

INTRODUCTION

A rapid worldwide increase has been observed regarding the prevalence of type 2 diabetes mellitus (DM) and obesity. The United Arab Emirates (UAE) is one of the epicentres of this epidemic. In a 2007 study, the agestandardised rates for pre-diabetes and type 2 DM among UAE nationals were 24.2% and 29%, respectively.¹ However, in a 2019 study

Strengths and limitations of this study

- This long-term longitudinal cohort study determines the incidence rate and risk factors of type 2 diabetes mellitus in overweight and obese United Arab Emirates (UAE) nationals.
- Anthropometric and laboratory data were obtained instead of self-reported information for the classification of risk factors and identification of incident cases.
- The study's subjects were recruited from a single large public hospital's ambulatory clinics; therefore, our findings may not apply to the general UAE population.
- No data were available for review regarding additional confounders, such as family history and changes in weight and physical activity.

of young Emirati men, the age-adjusted prevalence of pre-diabetes was much higher at 41.3%.² By contrast, the prevalence of type 2DM in North America and Western European countries is well below 12%.^{3–6} Notably, the number of people with type 2DM in the UAE is estimated to reach over two million by 2040.⁷

Overweight, defined as body mass index (BMI) between 25 and 29.99 kg/m^2 , and obesity, defined as BMI $\geq 30 \text{ kg/m}^2$, ⁸ are known risk factors of type 2DM.⁹ More recently, the age-adjusted prevalence of overweight and obesity among Emirati men between the ages of 18 and 30 years was noted to be 27% and 30%, respectively.² In contrast, the corresponding prevalence in another study was 23% and 10% among young Emirati women.¹⁰ Both these studies determined increased BMI to be strongly associated with diabetes.

Both obesity and type 2DM are well-known cardiometabolic risk factors for vascular complications, such as coronary heart disease

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Correspondence to Dr Saif Al-Shamsi; salshamsi@uaeu.ac.ae and peripheral vascular disease. The prevalence of these vascular diseases in the UAE is 10.5% and 11.1%, respectively.¹ These and other vascular-related complications have resulted in the UAE having one of the highest reported global death rates for cardiovascular disease—204 per 100 000 for women and 309 per 100 000 for men.¹¹

The rise in the incidence of diabetes and its complications among the UAE population has been in parallel to the economic surge, resulting in growing urbanisation and changing lifestyle habits in the country. Nevertheless, over the last decade, UAE has diverted its attention and endeavoured to curb the burden of overweight and obesity, as well as to decrease the incidence of associated conditions like diabetes. The Wegaya programme, a population-wide CVD screening programme, was initiated in 2008¹² to enable individuals and healthcare providers to implement CVD preventative measures. For continued assessment of the effectiveness of these local interventions, ad hoc analysis of population data at predefined intervals over a long period is required. However, epidemiological studies on type 2DM in the UAE have primarily focused on its prevalence in the general population.^{1 2 13} Despite having information from the Wegaya programme, data regarding the incidence of type 2DM among obese and overweight UAE nationals remain poorly researched. This lapse is primarily because of the significant resource investment required to conduct observational cohort studies with prolonged follow-up.

Therefore, in this long-term, longitudinal retrospective study, we aimed to calculate the incidence rate of type 2DM and elucidate the effects of BMI on DM risk in adult UAE nationals who are overweight and obese.

METHODS

Study setting

A retrospective electronic medical record review was performed of adult patients who presented to outpatient clinics at Tawam Hospital in Al Ain, UAE, between 1 April 2008 and 31 December 2008. Tawam Hospital is one of the two large government hospitals in the city of Al Ain and serves an estimated 60 000 adult UAE nationals in the city who are overweight and obese.^{14 15}

Subjects and procedures

The study population consisted of UAE nationals who were overweight and obese, aged between 18 and 75 years, who were seen at the outpatient departments at Tawam Hospital. The inclusion criteria were patients without type 2DM, defined as having glycated haemoglobin (HbA1c) <6.5%; patients not taking antidiabetic medications; and those with BMI \geq 25 kg/m² at baseline. The exclusion criteria were patients with missing HbA1c data, those with a history of chronic medical conditions, such as malignancy, HIV infection, receiving steroids or immunosuppression medications, or stage 3–5 chronic kidney disease (estimated glomerular filtration rate of \leq 59 mL/min/1.73 m²).¹⁶ Moreover, patients who had undergone

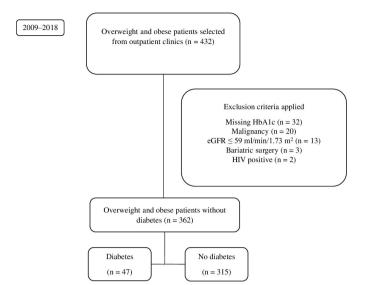


Figure 1 Flow chart for the inclusion and exclusion of patients. eGFR, estimated glomerular filtration rate; HbA1c, glycated haemoglobin.

weight reduction surgery, defined either as restrictive or malabsorptive bariatric surgery, or pharmacotherapy for weight loss were excluded from the study. The final study group consisted of 362 subjects with a BMI of $\geq 25 \text{ kg/m}^2$ (figure 1).

Definitions and measurements

Initial BMI was calculated from baseline height and weight measurements using the formula weight (kg) divided by height squared (m^2) . Overweight and obesity were defined according to the WHO classification,⁸ based on a BMI between 25 and 29.99 kg/m^2 and $\geq 30 \text{ kg/m}^2$, respectively. Hypertension (HTN) was defined as systolic blood pressure ≥140mm Hg, diastolic blood pressure \geq 90 mm Hg or use of blood pressure medications.¹⁷ The definition of dyslipidaemia was a serum total cholesterol (TC) of ≥6.21 mmol/L or if the patient had a documented prescription of lipid-lowering medications.¹⁸ Prediabetes was defined as having an HbA1c level between 5.7% and 6.4%.¹⁹ Current or history of smoking tobacco was considered a positive smoking history. Patients with a documented history of coronary heart disease, cerebrovascular disease or peripheral vascular disease were identified to have a vascular disease.

Outcomes

Our study endpoint was incident type 2DM, defined as either HbA1c \geq 6.5% (based on the definition of DM by the American Diabetes Association) or a diagnosis of type 2DM established by a physician at any time during the follow-up period.¹⁹ The occurrence of incident type 2DM was based on a review of all the electronic medical records until 30 April 2018. All laboratory analyses were performed at the Medical Laboratory Department of Tawam Hospital. An automated analyser, Integra 400 Plus (Roche Diagnostics, Mannheim, Germany), was used to measure HbA1c levels. The HbA1c results were reported as percentage using the International Federation of Clinical Chemistry-National Glycohemoglobin Standardization Program master equation.²⁰

Statistical analyses

Using a formula from Rosner,²¹ a sample size of 353 was calculated. This was based on the anticipated incidence of diabetes of $17.3\%^{22}$ and using 80% power at a two-sided significance level of 0.05. The sample size was further increased by 20% to account for patients lost to follow-up and those with missing data.

Intersex differences were determined using the Mann-Whitney U test or the independent-samples t-test for nonnormal and normal data, respectively. For categorical variables, Fisher's exact test (two-tailed) was used. Personyears at risk for incident type 2 DM was calculated for each patient as the time from the baseline visit to the diagnosis of type 2 DM or the most recent outpatient visit, whichever occurred first. The incidence rate of type 2 DM with a 95% CI was calculated per 1000 person-years by dividing the number of new cases of type 2 DM by the personyears at risk. Sex-specific incidence rates were calculated stratified by age and BMI categories. Kaplan-Meier timeto-event analysis was conducted with a log-rank test to compare age and BMI categories for incident type 2 DM.

Univariable and multivariable Cox proportional hazard models were evaluated to examine predictors of incident type 2DM in the entire cohort and stratified by sex. The following predictor variables were examined using univariable Cox modelling: age (categorised as <44 years and \geq 44 years), sex, vascular disease, HTN, dyslipidaemia, smoking, HbA1c (categorised as <5.7% and \geq 5.7%) and BMI (categorised as overweight and obese). All covariables with p<0.1 were included in the final multivariable model. A stepwise method based on likelihood ratios, with entry and removal probabilities set at 0.05 and 0.10, respectively, was used to check the results. The log-log plot was used to evaluate the proportional hazards assumption and was not significant. P<0.05 was considered statistically significant. SPSS V.25 was used to analyse the data.

Patient and public involvement

There was no patient or public involvement in the design and conduct of the study.

RESULTS

Characteristics of study participants

The baseline patient characteristics of the 362 subjects are presented in table 1. The ratio of men to women was almost 1:1. At baseline, the mean age was 44.4 ± 14.5 years and the BMI was 31.7 kg/m^2 (IQR 27.6–34.2 kg/m²). When stratified by sex, women were older, had higher BMI and TC values, and had a higher prevalence of dyslipidaemia at baseline. Conversely, men had significantly higher baseline systolic and diastolic blood pressure values, as well as a higher prevalence of smoking history.

Incidence

After a median follow-up time of 8.7 years (IQR, 8.2–9.0 years), 47 incident cases of type 2DM were identified.

Table 1 Baseline characteristics of the study participants									
	Total (N=362) Mean±SD	Men (n=170) Mean±SD	Women (n=192) Mean±SD	P value†					
Women (%)	53.0	-	-	_					
Age (years)	44.4±14.5	40.2±14.7	48.1±13.2	<0.001					
Anthropometric values									
BMI (kg/m ²)	31.7 (27.6–34.2)	29.6 (27.3–32.8)	31.2 (27.9–35.8)	0.001					
SBP (mm Hg)	127.9±17.3	129.9±17.9	126.2±16.7	0.040					
DBP (mm Hg)	76.9±11.0	79.0±11.3	75.1±10.4	0.001					
Laboratory values									
HbA1c (%)	5.6±0.5	5.6±0.5	5.6±0.5	0.326					
eGFR (mL/min/1.73 m ²)	105.9±16.8 (n=359)*	104.2±18.5 (n=167)*	107.4±15.1	0.073					
TC (mmol/L)	5.1±0.9	4.9±1.0	5.2±1.0	0.002					
History of, n (%)									
HTN	181 (50.0)	91 (53.5)	90 (46.9)	0.247					
Dyslipidaemia	141 (39.0)	52 (30.6)	89 (46.4)	0.002					
Vascular disease	18 (5.0)	10 (5.9)	8 (4.2)	0.478					
Smoking	56 (15.5)	54 (31.8)	2 (1.0)	<0.001					

*The difference in n is due to missing data.

†Independent-samples t-test was used to calculate p values for continuous variables and Fisher's exact test (two-tailed) for categorical variables.

BMI, body mass index; DBP, diastolic blood pressure; eGFR, estimated glomerular filtration rate; HbA1c, glycated haemoglobin; HTN, hypertension; SBP, systolic blood pressure; TC, total cholesterol.

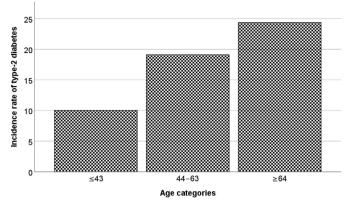


Figure 2 Overall incidence rate of type 2 diabetes mellitus (cases/1000 person-years) by increasing age.

The overall incidence rate of type 2DM was 16.3 (95% CI 12.1 to 21.4) cases per 1000 person-years, and the crude incidence over the same period was 13.0% (95% CI 9.8 to 16.8%). The incidence rate of type 2DM increased with patient age (figure 2). When stratified by sex, the incidence rates were 17.7 (95% CI 11.6 to 25.9) and 15.0 (95% CI 9.8 to 22.2) cases per 1000 person-years in men and women, respectively.

Sex-specific incidence rates, as stratified by BMI, were higher in obese women (21.1, 95% CI 13.3 to 32.0 cases per 1000 person-years) compared with overweight women (5.1, 95% CI 1.3 to 14.0 cases per 1000 person-years) (p=0.013). Incidence rates for type 2DM were higher in older women (21.3, 95% CI 13.5 to 31.9 cases per 1000 person-years) compared with younger women (3.7, 95% CI 0.6 to 12.2 cases per 1000 person-years) (p=0.008) (figure 3). However, these differences were not significant among men.

Analyses of risk factors

We used multivariable Cox regression analyses to examine the association of age, sex, vascular disease, dyslipidaemia, smoking, HTN, BMI and HbA1c levels with the risk of developing type 2 DM in the entire study cohort, as well as between men and women. When adjusted for all other variables, we observed that in men an HbA1c level \geq 5.7% (HR=3.02, 95% CI 1.20 to 7.62) and in women age \geq 44 years (HR=4.84, 95% CI 1.13 to 20.73) and obesity (HR=3.42, 95% CI 1.01 to 11.57) were significant risk factors for the development of type 2 DM (table 2).

DISCUSSION

The results of this study revealed that the crude incidence of type 2DM among UAE nationals who are overweight and obese was 1.49% per year during the 9-year study period. The overall incidence rate was 16.3 per 1000 person-years. The independent predictors for the development of type 2DM were age (≥44 years) and obesity in women and pre-diabetes in men.

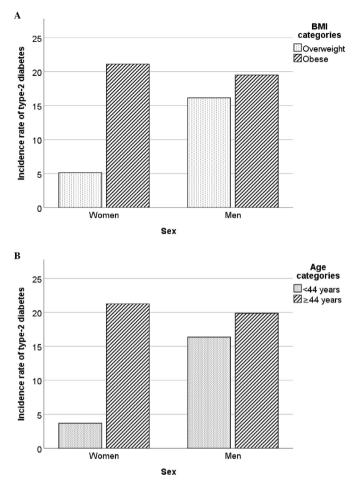


Figure 3 Sex-specific incidence rate of type 2 diabetes mellitus (cases/1000 person-years) stratified by BMI categories (A) and by age (B). BMI, body mass index.

Incidence rates

The incidence rates of type 2DM in high-risk populations differ worldwide. This difference is probably due to variations in insulin sensitivity related to the interaction between genetic and environmental factors among different ethnic and population groups globally.²³ In a US study of predominantly overweight and obese African-American women, the reported incidence rate of type 2DM was 15.2 per 1000 person-years²⁴ and is comparable with our study. There has been only one other study that has assessed the incidence rate of type 2 DM in the UAE.¹³ The study was conducted over 12 months in Ajman, one of the emirates in the UAE, and revealed that the incidence rate for type 2DM was 4.8 per 1000 person-years. However, this study cannot be directly compared with our study because it was conducted on the general population, and our study cohort comprised only overweight and obese UAE nationals.

Risk factors

This study revealed some interesting findings regarding the risk factors for developing type 2DM in the highrisk UAE population. Sex-specific incidence rates of type 2DM exhibited a female preponderance, particularly older women. Nevertheless, these results were not

	Total (N=362)		Men (n=170)		Women (n=192)	
Predictor variables	HR (95% CI)	P value	HR (95% CI)	P value	HR (95% CI)	P value
Age						
<44 years	Reference		Reference		Reference	
≥44 years	1.20 (0.59 to 2.44)	0.619	0.46 (0.17 to 1.13)	0.135	4.84 (1.13 to 20.73)	0.034
Sex						
Female	Reference					
Male	1.60 (0.83 to 3.05)	0.160				
History of HTN						
No	Reference		Reference		Reference	
Yes	1.78 (0.94 to 3.37)	0.075	1.89 (0.74 to 4.81)	0.183	1.61 (0.67 to 3.88)	0.292
History of dyslipidaemia						
No	Reference		Reference		Reference	
Yes	1.32 (0.70 to 2.51)	0.393	2.01 (0.80 to 5.04)	0.135	0.80 (0.34 to 1.90)	0.608
History of smoking						
No	Reference		Reference		Reference	
Yes	0.58 (0.23 to 1.49)	0.259	0.36 (0.12 to 1.04)	0.060	4.88 (0.44 to 54.08)	0.196
History of vascular disease						
No	Reference		Reference		Reference	
Yes	1.57 (0.54 to 4.54)	0.408	2.46 (0.65 to 9.27)	0.183	0.60 (0.06 to 5.50)	0.647
HbA1c						
<5.7%	Reference		Reference		Reference	
≥5.7%	2.29 (1.16 to 4.52)	0.016	3.02 (1.20 to 7.62)	0.019	2.30 (0.83 to 6.39)	0.109
BMI						
Overweight	Reference		Reference		Reference	
Obese	1.86 (0.99 to 3.50)	0.053	1.17 (0.51 to 2.68)	0.713	3.42 (1.01 to 11.57)	0.048

BMI, body mass index; HbA1c, glycated haemoglobin; HTN, hypertension.

unique and were also observed among the general population of the UAE.¹³ The higher incidence of type 2DM among older women may be attributable to the high prevalence of overweight and obesity, sedentary lifestyle, nutritional habits and early menopause witnessed in our population.^{25–28}

Furthermore, individuals with high-normal HbA1c levels have a greater risk of developing type 2DM.²⁹ However, when we stratified our cohort by sex, the independent predictive property of the HbA1c level was only observed in men. An explanation for this finding could be the high prevalence of smoking history in men, as smoking has been shown to increase HbA1c levels in individuals without diabetes.³⁰ Moreover, obesity in women was noted to be an independent risk predictor for type 2DM. Studies have demonstrated that the risk of developing type 2DM is significantly higher in individuals with BMI $\geq 30 \text{ kg/m}^2$ compared with those with normal BMI.⁹ Therefore, interventions that promote weight loss and reduce BMI can reduce the risk of developing type 2DM. Notably, an 8-week exercise intervention programme in a UAE student population demonstrated a significant

reduction in BMI.³¹ In addition, other controlled intervention studies in high-risk populations^{32,33} demonstrated a significant decrease in the risk of type 2DM consequent to BMI reduction.

Clinical and public health implications

In the UAE, much of the increase in type 2DM has occurred due to social and economic changes—from traditional, seminomadic lifestyle to sedentary, urban and high-income lifestyle. Consequently, the progression towards type 2DM and cardiovascular disease starts early in the UAE due to the high prevalence of childhood obesity.^{1 34} A 2015 study of Emirati children and adolescents with a BMI \geq 25 kg/m² determined the prevalence of pre-diabetes and type 2DM to be 5.4% and 0.87%, respectively.³⁵ Therefore, reducing the burden of type 2DM in the UAE will require intensive population-wide interventions to promote regular physical activity and a healthy diet,^{33 36} particularly among the Emirati youth.

Furthermore, the results of this study highlight the need to focus on Emirati women as a specific target group for obesity prevention programmes. These programmes should begin in childhood because 21% are obese by the time they are 30 years of age.²⁶ In addition, it is imperative to have a multidisciplinary approach to address the high incidence of type 2DM, where paediatricians, primary care providers, dietitians and specialists work collaboratively towards promoting weight reduction and maintaining a healthy BMI in our population.

Strengths and limitations

The strengths of this study include the long duration of follow-up for our representative sample of overweight and obese individuals in the second largest city in the Abu Dhabi Emirate of UAE. We used the anthropometric and laboratory data documented in patient charts instead of self-reported information and diagnostic codes for the classification of risk factors and identification of incident cases. In addition, HbA1c levels or physician diagnosis formed the basis to identify incident cases of type 2DM instead of fasting plasma glucose, which has more variability.^{37 38}

Nonetheless, this study had some limitations. Baseline data regarding family history of type 2DM, history of gestational diabetes, physical activity and abdominal obesity were unavailable—all of which are well-researched risk factors for the development of DM. Nevertheless, a prospective study can provide more robust results. Furthermore, data on additional confounders, such as family history, changes in weight and physical activity, were unavailable for review. Finally, the subjects for this study were recruited from a single largest tertiary care hospital's ambulatory clinics; thus, our findings may not apply to the general UAE population.

CONCLUSIONS

The overall incidence rate of type 2 DM in UAE nationals who are overweight and obese was 16.3 per 1000 personyears. Notably, pre-diabetes in men and obesity in women, particularly older women, were determined to be independent predictors for the development of incident type 2 DM. Hence, it is imperative to emphasise BMI reduction through lifestyle modifications in these high-risk groups.

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Contributors DR conceived the study and its design, performed the literature search and drafted the initial manuscript. SA-S was involved in data management, statistical analysis, interpretation of data and critical review of the manuscript. RDG assisted in statistical analysis, contributed to the discussion, and reviewed and edited the manuscript. JA-K performed a critical review of the manuscript, was involved in data interpretation and also guided the research methodology. All authors read and approved the final manuscript.

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Competing interests None declared.

Patient consent for publication Not required.

Ethics approval The Research and Ethics Board of Tawam Hospital and the United Arab Emirates University granted ethical approval to this study (CRD239/13).

Informed consent was waived because patient records and information were anonymised and deidentified before analysis.

Provenance and peer review Not commissioned; externally peer reviewed.

Data availability statement The data that support the findings of this study are available from the corresponding author, SA-S, upon reasonable request.

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