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# **BMJ Open**

# Incidence and risk factors of type-2 diabetes mellitus in an overweight and obese population: a long-term study from a Gulf State

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5 6	2	population: a long-term study from a Gulf State
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10	4	Dybesh Regmi <sup>1</sup> Saif Al-Shamsi <sup>2*</sup> Romona D. Govender <sup>3</sup> Juma Al Kaabi <sup>2</sup>
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2 3 4	24	Abstract
5 6	25	<b>Objectives:</b> A high body mass index (BMI) is associated with a risk for type-2 diabetes mellitus
7 8 9	26	(DM). The United Arab Emirates (UAE) is experiencing a marked increase in obesity. Data on
10 11	27	the incidence of type-2 DM in this high-risk adult UAE population is unavailable. This study
12 13	28	aimed to evaluate the incidence and risk of developing type-2 DM among individuals with
14 15 16	29	above-normal BMI in the UAE.
17 18	30	Design: A retrospective cohort study.
19 20	31	Setting: Outpatient clinics at a tertiary care centre in Al Ain, UAE.
21 22 23	32	Participants: 362 overweight or obese adult UAE nationals who visited outpatient clinics
23 24 25	33	between April 2008 and December 2008.
26 27	34	Primary outcome measure: Patients with type-2 DM were identified based on diagnosis
28 29	35	established by a physician or glycated hemoglobin (HbA1c) levels $\geq 6.5\%$ during the follow-up
30 31 32	36	period (until April 2018).
33 34	37	<b>Results:</b> The overall incidence rate of type-2 DM during the median follow-up time of 8.7 years
35 36	38	was 16.3 (95% CI: 12.1–21.4) cases per 1000 person-years. Incidence rates in males and females
37 38 39	39	were 17.7 (95% CI: 1.14–2.60) and 15.0 (95% CI: 0.99–2.35), respectively. Using multivariable
40 41	40	Cox proportional hazard analysis, older age and obesity in females and HbA1c levels in males
42 43	41	were determined to be independent risk factors for developing type-2 DM.
44 45 46	42	<b>Conclusions:</b> The incidence rate of type-2 DM in overweight and obese UAE nationals is high.
40 47 48	43	In addition to screening, current strategies should strongly emphasize lifestyle modifications that
49 50	44	decrease HbA1c and BMI levels in this high-risk population.
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#### Strengths and limitations of this study

A long-term (9-years) study that utilizes anthropometric and laboratory data to accurately quantify previously unknown incidence rate of type-2 DM in a high-risk group in the United Arab Emirates.

- In doing so, the results demonstrate different independent risk factors that influence the • development of type-2 DM in males and females suggesting unique focused gender based strategies to be incorporated into the current preventative program.
- The study results provide a baseline to assess effectiveness of national diabetes preventative programs in obese and overweight individuals while allowing epidemiologists to benchmark the incidence rates globally.
- The usual limitations of a retrospective study apply here as well. A prospective study would probably also allow capturing data on physical activity, and changes in weight, both of which can influence development of type-2 DM.
- Another limitation of this study is missing baseline data on family history and history of
  - Gestational diabetes both of which are well-known risk factors for developing type 2-DM.
- Their contribution if any to the development of new onset type-2 DM in our study population is not accounted for.

Keywords: incidence; obesity; risk factors; type-2 diabetes mellitus; United Arab Emirates

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**INTRODUCTION** There has been a rapid increase in the prevalence of obesity and type-2 diabetes mellitus (DM) globally. The United Arab Emirates (UAE) is one of the epicenters of this epidemic. Among UAE nationals, the crude prevalence of obesity and type-2 DM is 37%[1] and 29%[2], respectively and the number of people with type-2 DM in the country is estimated to reach over 2 million by 2040.[3] Obesity, defined as a body mass index (BMI)  $\geq$  30 kg/m[2,4], is a known risk factor for type-2 DM<sup>5</sup> while lifestyle interventions such as weight loss in obese individuals with impaired glucose tolerance have been shown to reduce the risk for type-2 DM.[6] There has been significant attention and effort over the last decade in the UAE to curb the prevalence of overweight and obesity, as well as to decrease the incidence of associated diseases. The Wegava, a population-wide CVD screening program, was initiated in 2008[7] to enable individuals and healthcare providers to implement preventative measures. Epidemiological studies on type-2 DM in the UAE have primarily focused on its prevalence in the general population.[8] A decade after the implementation of the preventive program, data on the incidence of type-2 DM among obese and overweight UAE nationals remain unknown. In this retrospective longitudinal cohort study, we aimed to determine the incidence of DM and elucidate the effect of BMI on type-2 DM risk in overweight and obese adult UAE nationals. **METHODS Study setting** A retrospective electronic medical record review of adult patients who presented to outpatient clinics at Tawam hospital in Al Ain, UAE between April 1st, and December 31st 2008 was 

performed. The tertiary care hospital is one of two governmental hospitals in the city and serves
an estimated population of 650,000 patients, of which 30% are UAE nationals.[9] Tawam
Hospital and the UAE University research and ethics board granted ethical approval for this
study (CRD239/13). Informed consent was waived because patient records and information were
anonymized and de-identified prior to analysis.

99 Subjects and procedures

The study population consisted of healthy overweight and obese UAE nationals, aged between 18 and 75 years, who were seen at primary care and specialty outpatient clinics at Tawam hospital (obstetrics and gynecology, internal medicine, and surgery, as well as subspecialty clinics). The inclusion criteria were patients without type-2 DM, defined as having glycated hemoglobin (HbA1c) <6.5%; patients not taking anti-diabetic medications; and those with BMI  $\geq$  25 kg/m<sup>2</sup> at baseline. The exclusion criteria were patients with missing HbA1c data; those with a history of chronic medical conditions such as malignancy, human immunodeficiency virus infection, receiving steroids or immunosuppression medications, or stage 3–5 chronic kidney disease [estimated glomerular filtration rate (eGFR) of  $\leq$ 59 mL/min/1.73 m<sup>2</sup>][10]. Those who had undergone weight reduction surgery defined as either restrictive or malabsorptive bariatric surgery or pharmacotherapy for weight loss were also excluded from the study. The final study group consisted of 362 subjects with a BMI of  $\geq 25 \text{ kg/m}^2$  (Fig 1). 

- 7 112
  - 113 Definitions and measurements

Initial BMI was calculated from height and weight measured at baseline using the formula
weight (kg) divided by height squared (m<sup>2</sup>). Overweight and obesity were defined according to

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the World Health Organization classification[4], with a BMI between 25 and 29.99 kg/m<sup>2</sup> and  $\geq$ 30 kg/m<sup>2</sup>, respectively. Hypertension (HTN) was defined as systolic blood pressure  $\geq$ 140 mmHg, diastolic blood pressure  $\geq$ 90 mmHg, or the use of antihypertensive medications.[11] Dyslipidemia was defined as serum total cholesterol (TC) of  $\geq 6.21$  mmol/L or if the patient had a documented prescription of lipid-lowering medications.[12] Current or prior history of smoking tobacco was considered positive for smoking history. Patients were considered to have vascular disease if they had a documented history of either coronary heart disease, cerebrovascular disease, or peripheral vascular disease. Outcomes Our study endpoint was incident type-2 DM, defined as either HbA1c  $\geq$ 6.5% or a diagnosis of type-2 DM established by a physician occurring at any time during the follow-up period.[13] The occurrence of incident type-2 DM was based on a review of all of the electronic medical records until April 30, 2018. All laboratory assays at baseline and follow-up were conducted at the Medical Laboratory Department at Tawam Hospital. HbA1c levels were measured using an automated analyzer Integra 400 Plus (Roche Diagnostics, Mannheim, Germany). **Statistical analyses** Differences between sexes were determined using an independent-samples t-test or the Mann-Whitney U test for normal and non-normal data, respectively, with Fisher's exact test (two-tailed) being used for categorical variables. Person-years at risk for developing type-2 DM was calculated for each subject as the time from 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 95% 

1 2		
2 3 4	139	confidence intervals (CI) was calculated per 1000 person-years by dividing the number of new
5 6	140	cases of type-2 DM by the person-years at risk. Sex-specific incidence rates stratified by age and
/ 8 9	141	BMI categories were calculated. Kaplan-Meier time-to-event analysis was conducted with a log-
9 10 11	142	rank test to compare age and BMI categories for incident type-2 DM.
12 13	143	Univariable and multivariable Cox proportional hazards modeling (stratified by sex) were
14 15 16	144	used to examine predictors of incident type-2 DM. The following predictor variables were
17 18	145	examined by univariable Cox modeling: age (categorized as <44 years and $\geq$ 44 years), sex
19 20	146	(categorized as female/male), vascular disease (categorized as yes/no), HTN (categorized as
21 22 22	147	yes/no), dyslipidemia (categorized as yes/no), smoking (categorized as yes/no), HbA1c
23 24 25	148	(categorized as $<5.7\%$ and $\ge 5.7\%$ ), and BMI (categorized as overweight and obese). All
26 27	149	covariables with P <0.1 were included in the final multivariable modeling. Results were checked
28 29 30 31 32 33 34	150	by a step-wise method based on likelihood ratios, with entry and removal probabilities set at 0.05
	151	and 0.10, respectively. The proportional hazards assumption was tested using a log-log plot and
	152	were not significant. A P-value < 0.05 was considered statistically significant. SPSS version 25
35 36	153	(IBM Corp., Armonk, NY) was used to analyze the data.
37 38 20	154	
40 41	155	Patient and public involvement
42 43	156	There was no patient or public involvement in the design and conduct of the study.
44 45	157	
40 47 48	158	RESULTS
49 50	159	Characteristics of study participants
51 52	160	The baseline patient characteristics, including demographic, clinical, and biochemical data of the
53 54 55	161	362 subjects, are presented in Table 1. The ratio of males to females was almost 1:1. The mean
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age and BMI at baseline were  $44.36 \pm 14.45$  years and 31.69 [interquartile range (IQR) 27.64, 34.18 kg/m<sup>2</sup>], respectively. When stratified by sex, females were older, had higher BMI and TC values, and had a higher prevalence of dyslipidemia, at baseline. Conversely, males had significantly higher baseline systolic and diastolic blood pressure values, as well as a higher prevalence of smoking history.

168 **Table 1. Baseline characteristics of the study participants.** 

			Females		
	Total ( $N=362$ )	Males $(N=170)$	(N=192)	p-value*	
	Mean ± SD	Mean $\pm$ SD	Mean ± SD	-	
Females (%)	53.0	-	-	-	
Age (years)	$44.36 \pm 14.45$	$40.2 \pm 14.7$	$48.09 \pm 13.2$	< 0.001	
Anthropometric values					
$\mathbf{DMI}\left(1_{ra}/m^{2}\right)$	31.69 (27.64,	29.56 (27.33,	31.22 (27.91,	0.001	
Divii (kg/iii-)	34.18)	32.79)	35.77)	0.001	
SBP (mmHg)	$127.93 \pm 17.34$	$129.92 \pm 17.86$	$126.17 \pm 16.72$	0.040	
DBP (mmHg)	$76.94 \pm 11.01$	79.03 ± 11.34	$75.09 \pm 10.40$	0.001	
Laboratory values		0			
HbA1c (%)	$5.62 \pm 0.48$	5.59 ± 0.50	$5.64 \pm 0.46$	0.326	
$eGFR (mI/min/1.73 m^2)$	$105.87 \pm 16.81$	104.16 ± 18.47	$107.35 \pm 15.13$	0.073	
	(N=359)†	(N=167)†	107.55 ± 15.15	0.075	
TC (mmol/L)	$5.06 \pm 0.88$	$4.89 \pm 0.95$	$5.21 \pm 0.99$	0.002	
History of n (%)					
HTN	181 (50.0)	91 (53.5)	90 (46.9)	0.247	
Dyslipidemia	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	

1 2		
- 3 4	169	HTN hypertension, BMI body mass index, eGFR estimated glomerular filtration rate, SBP
5 6 7 8 9 10 11 12 13 14 15	170	systolic blood pressure, DBP diastolic blood pressure, SD standard deviation, HbA1c glycated
	171	hemoglobin, TC total cholesterol.
	172	†The difference in N is due to missing data.
	173	*Independent-samples t-test was used to calculate p values for continuous variables and Fisher's
	174	exact test (two-tailed) for categorical variables.
16 17	175	
18 19 20	176	Incidence
21 22	177	After a median follow-up time of 8.7 years (IQR, 8.2–9.0 years), 47 incident cases of type-2 DM
23 24	178	were identified. The overall incidence rate of type-2 DM was 16.3 (95% CI: 12.1-21.4) cases per
25 26 27 28 29 30 31	179	1000 person-years and the crude incidence of type-2 DM over the same period was 13.0% (95%
	180	CI: 9.8–16.8%).
	181	Sex-specific incidence rates, as stratified by BMI, were higher in obese females (21.1, 95%
32 33 34	182	CI: 13.3-32.0 cases per 1000 person-years), when compared to overweight females (5.1, 95%
35 36	183	CI: 1.3-14.0 cases per 1000 person-years) (P=0.013). Incidence rates for type-2 DM were higher
37 38	184	in older females (21.3, 95% CI: 13.5-31.9 cases per 1000 person-years), when compared to
39 40 41	185	younger females (3.7, 95% CI: 0.6–12.2 cases per 1000 person-years) (P=0.008), (Figs. 2 and 3,
42 43	186	respectively). However, these differences were not significant among males.
44 45	187	Analyses of risk factors
46 47 48 49	188	We used multivariable Cox regression analyses to examine the association of age, vascular
	189	disease, dyslipidemia, smoking, HTN, BMI, and HbA1c levels with the risk of developing type-2
51 52	190	DM between males and females. When adjusted for all other variables, we found that in males,
53 54 55 56 57	191	an HbA1c level $\geq$ 5.7% [hazard ratio (HR) = 3.02, 95% CI: 1.20–7.62] and for females age $\geq$ 44

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192	years (HR = 4.84, 95% CI:	1.13-20.73) and obe	esity (HR = 3	6.42, 95% CI: 1.01–1	1.57) were
193	significant risk factors for the	he development of ty	vpe-2 DM (T	able 2).	
194					
195	Table 2. Predictors of inci	dent type-2 diabete	s mellitus in	the study population	on.
	Predictor variables	Male (N=170)		Female (N=192)	
		HR (95% CI)	p-value	HR (95% CI)	p-value
	Age				
	<44 years	Reference		Reference	
	≥44 years	0.46 (0.17–1.13)	0.135	4.84 (1.13-20.73)	0.034
	History of HTN				
	No	Reference		Reference	
	Yes	1 89 (0 74-4 81)	0.183	1 61 (0 67–3 88)	0.292
	History of dyslinidemia		0.105	1.01 (0.07 5.00)	0.272
	No	Pafaranaa		Poforonoo	
	No.	$\begin{array}{c} \text{Reference} \\ 2.01 (0.80, 5.04) \end{array}$	0.125	$\begin{array}{c} \text{Reference} \\ 0.90 (0.24, 1.00) \end{array}$	0.609
	<u>1 es</u>	2.01 (0.80-3.04)	0.133	0.80 (0.34–1.90)	0.008
	History of smoking	D.C.		D.C.	
	NO	Reference	0.000	Reference	0.106
	Yes	0.36 (0.12–1.04)	0.060	4.88 (0.44–54.08)	0.196
	History of vascular				
	disease				
	No	Reference		Reference	
	Yes	2.46 (0.65–9.27)	0.183	0.60 (0.06–5.50)	0.647
	HbA1c				
	<5.7%	Reference		Reference	
	≥5.7%	3.02 (1.20-7.62)	0.019	2.30 (0.83-6.39)	0.109
	BMI				
	Overweight	Reference		Reference	
	Obese	1 17 (0 51-2 68)	0.713	3 42 (1 01–11 57)	0.048
196	HTN hypertension, BMI boo	dy mass index, <i>HbA</i>	<i>c</i> glycated h	emoglobin, <i>CI</i> confic	lence intervals,
197	HR hazard ratio.				
198					
199	DISCUSSION				
200	The results of this study sho	owed that the inciden	ice of type-2	DM among overweig	ght and obese
201	UAE nationals was 1.49% p	per year during the 9	-year study p	period. The overall in	cidence rate
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was 16.3 per 1000 person-years. The independent risk factors for the development of type-2 DM were age ( $\geq$  44 years) and obesity in females and HbA1c levels in males. 

#### **Incidence** rates

The incidence rates of type-2 DM in the general population differ significantly worldwide. This is possibly due to variations in insulin sensitivity related to the interaction between genetic and environmental factors among different populations globally.[14] A study conducted over a 12month period in Ajman, one of the emirates in the UAE, the incidence rate for type-2 DM in the general Emirati population was 4.8 per 1000 person-years[8], while our study had an incidence rate 3 times higher. This is not unexpected as our study focused on overweight and obese individuals and is comparable to other research in high-risk populations in the United States (15.2 per 1000 person-years)[15], while South-Asian countries e.g. India and Bangladesh reported incidence rates of 22.2 per 1000 person-years [16] and 16.4 per 1000 person-years L.C. respectively.[17] 

#### **Risk factors**

This study revealed some interesting findings regarding the risk factors for developing type-2 DM in this high-risk UAE population. Sex-specific incidence rates of type-2 DM showed a female preponderance, particularly among older females. These results were not unique and were also observed among the general population of the UAE.[8] The higher incidence of type-2 DM among older females may be attributable to the high prevalence of overweight and obesity [18], sedentary lifestyle, nutritional habits[19], and early menopause witnessed in our population.[20] Local data shows that for every 1-year increase in age, the risk of obesity among females increases by 5%.[21]

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Individuals with high-normal HbA1c levels have a greater risk of developing type-2 DM.[22] However, in our study, the independent predictive property of the HbA1c level was only observed in males. An explanation for this finding could be the high prevalence of male smokers seen, as smoking has been shown to increase HbA1c levels in people without diabetes.[23] Obesity in females was also an independent risk predictor for type-2 DM. Studies have shown that the risk of developing type-2 DM is significantly higher in individuals with BMI  $\geq$  30 kg/m<sup>2</sup> when compared to those with normal BMI.[5] Interventions that promote weight loss and decreased BMI, also reduce the risk of developing type-2 DM. An 8-week exercise intervention program in a UAE student population demonstrated a significant reduction in BMI.[24] Other controlled intervention studies in high-risk populations [6,25,26] also demonstrated a significant decrease in the risk of type-2 DM as a result of BMI reduction. 

# **Clinical and public health implications**

Results from this study demonstrate that the incidence of type-2 DM in the high-risk UAE population begins to increase by the fourth decade of life, and this trend significantly increases with higher BMI values. Thus, targeting overweight or obesity, which are modifiable risk factors for type-2 DM[27], is crucial in reducing this burden. A 2015 study of Emirati adolescents found the prevalence of pre-diabetes and type-2 DM in overweight and obese children were 5.8% and 0.87%, respectively.[28] Although alarming, the results are unsurprising when we consider the high prevalence of childhood obesity in the UAE. Furthermore, the rates of overweight and obesity in young UAE adults was 36%[29], putting them at risk for developing type-2 DM. Screening programs alone are inadequate and clinicians must strongly recommend lifestyle modifications to reduce overweight and obesity.

**Strengths and limitations** 

The strengths of this study include the duration of follow-up for our representative sample of overweight and obese UAE nationals in the second-largest city in the Abu Dhabi Emirate of UAE. We utilized measured anthropometric and laboratory data in patient charts instead of selfreported information and diagnostic codes for the classification of risk factors and identification of incident cases. In addition, HbA1c levels or documented diagnosis by a physician were used to identify incident cases of type-2 DM instead of fasting plasma glucose, which has more variability.[30,31] This study has some limitations. These limitations include missing baseline data regarding family history of type-2 DM, history of gestational diabetes, physical activity, and abdominal 

obesity, all of which are well-researched risk factors for the development of DM. A prospective study will ensure more robust data, as well as changes in lifestyle variables such as exercise, 

weight loss or gain. 

#### CONCLUSIONS

The overall incidence rate of type-2 DM in overweight and obese UAE nationals was 16.3 per 1000 person-years. The HbA1c level in males and obesity in females, particularly in the older age group, were found to be independent risk factors for the development of type-2 DM. Thus, there should be a greater emphasis on reducing BMI through lifestyle modifications in these high-risk groups.

#### Acknowledgments

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1		
2 3 4	269	Author Contributions D.R. conceived the study and its design, performed literature search and
5 6	270	drafted the initial manuscript. S.A.S. was involved in data management, statistical analysis,
/ 8 9	271	interpretation of the data, and critical review of the manuscript. R.D.G. assisted in statistical
10 11	272	analysis, contributed to the discussion, reviewed and edited the manuscript. J.A.K. performed a
12 13	273	critical review of the manuscript was involved in data interpretation and also provided guidance
14 15 16	274	in research methodology. All authors read and approved the final manuscript.
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24 25	278	decision to publish, or preparation of the manuscript.
26 27	279	Competing interests
28 29 30	280	The authors declare that they have no competing interests.
31 32	281	Patient Consent for publication
33 34	282	Not required
35 36 27	283	Availability of data and materials
37 38 39	284	The dataset used and analyzed during the current study is available from the corresponding
40 41	285	author on reasonable request.
42 43	286	
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36 37 38	384		1545–1551.			
39 40	9 <sup>0</sup> 385 FIGURE LEGENDS					
41 42	505	ПС				
43 44	386	Fig 1. Flow chart for the inclusion and exclusion of patients. HbA1c glycated hemoglobin,				
45 46	387	eGF	eGFR estimated glomerular filtration rate, HIV human immunodeficiency virus.			
47 48	388	Fig	2. The sex-specific incidence rate of type-2 diabetes mellitus (cases/1000 person-years)			
49 50	389	stratified by BMI categories. BMI body mass index.				
51 52 53	390	3. The sex-specific incidence rate of type-2 diabetes mellitus (cases/1000 person-years)				
54 55	391	stratified by age.				
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Males

BMI

categories

⊡Overweight ⊠Obese







The sex-specific incidence rate of type-2 diabetes mellitus (cases/1000 person-years) stratified by age.

144x84mm (300 x 300 DPI)

STROBE 2007 (v4) Statement—Checklist of items that should be included in reports of <i>cohort studies</i>			
Section/Topic	Item #	Recommendation	Reported on page #
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract	2
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	2
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	4
Objectives	3	State specific objectives, including any prespecified hypotheses	4
Methods			
Study design	4	Present key elements of study design early in the paper	4, 5
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	
Participants	6	(a) Give the eligibility criteria, and the sources and methods of selection of participants. Describe methods of follow-up	5
		(b) For matched studies, give matching criteria and number of exposed and unexposed	NA
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable 5,6	
Data sources/ measurement	Data sources/       8*       For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe       6,7         neasurement       comparability of assessment methods if there is more than one group       comparability of assessment methods if there is more than one group		6,7
Bias	9	Describe any efforts to address potential sources of bias	
Study size	10	Explain how the study size was arrived at	5
Quantitative variables       11       Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why       6,7		6,7	
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	7
		(b) Describe any methods used to examine subgroups and interactions	7
		(c) Explain how missing data were addressed	NA
		(d) If applicable, explain how loss to follow-up was addressed	NA
		(e) Describe any sensitivity analyses	NA
Results			

Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility, confirmed	5
		eligible, included in the study, completing follow-up, and analysed	
		(b) Give reasons for non-participation at each stage	5
		(c) Consider use of a flow diagram	Figure 1
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential	7,8
		confounders	
		(b) Indicate number of participants with missing data for each variable of interest	8
		(c) Summarise follow-up time (eg, average and total amount)	9
Outcome data	15*	Report numbers of outcome events or summary measures over time	9
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence	9,10
		interval). Make clear which confounders were adjusted for and why they were included	
		(b) Report category boundaries when continuous variables were categorized	9,10
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	NA
Other analyses 17 Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses		9,10	
Discussion			
Key results	18	Summarise key results with reference to study objectives	10
Limitations			
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from	11,12
		similar studies, and other relevant evidence	
Generalisability	21	Discuss the generalisability (external validity) of the study results	11
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on	13,14

\*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

**Note:** An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.

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# **BMJ Open**

## Incidence and risk factors of type 2 diabetes mellitus in an overweight and obese population: a long-term retrospective study from a Gulf State

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Complete List of Authors:	Regmi, Dybesh; University of Toronto Faculty of Medicine, Department of Family and Community Medicine Al-Shamsi, Saif; UAE University College of Medicine and Health Sciences, Department of Internal Medicine Govender, Romona; UAE University College of Medicine and Health Sciences, Department of Family Medicine Al-Kaabi, Juma; UAE University College of Medicine and Health Sciences, Department of Internal Medicine
<b>Primary Subject Heading</b> :	Diabetes and endocrinology
Secondary Subject Heading:	Epidemiology
Keywords:	DIABETES & ENDOCRINOLOGY, EPIDEMIOLOGY, INTERNAL MEDICINE





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2 3	1	Incidence and risk factors of type 2 diabetes mellitus in an overweight and obese		
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6	2	population: a long-term retrospective study from a Gulf State		
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10 11	4	Dybesh Regmi <sup>1</sup> , Saif Al-Shamsi <sup>2*</sup> , Romona D. Govender <sup>3</sup> , Juma Al Kaabi <sup>2</sup>		
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2	24	Abstract				
4	24	ADSIFACI				
5 6 7	25	<b>Objectives:</b> A high body mass index (BMI) is associated with a risk of type 2 diabetes mellit				
7 8 9	26	(DM). The United Arab Emirates (UAE) is experiencing a marked increase in obesity.				
10 11	27	Nonetheless, no data is available regarding the incidence of type 2 DM in the high-risk adult				
12 13	28	UAE population. Therefore, this study aimed to evaluate the incidence rate and risk of				
14 15 16	29	developing type 2 DM among individuals with above-normal BMI in the UAE.				
16 17 18	30	Design: A retrospective cohort study.				
19 20	31	Setting: Outpatient clinics at a tertiary care center in Al Ain, UAE.				
21 22	32	Participants: Three hundred sixty-two overweight or obese adult UAE nationals who visited				
23 24 25	33	outpatient clinics between April 2008 and December 2008.				
25 26 27	34	Primary outcome measure: Patients with type 2 DM were identified based on diagnosis				
28 29	35	established by a physician or through glycated hemoglobin (HbA1c) levels $\geq 6.5\%$ during the				
30 31 32	36	follow-up period (until April 2018).				
33 34	37	<b>Results:</b> The overall incidence rate of type 2 DM during the median follow-up time of 8.7 years				
35 36	38	was 16.3 [95% confidence interval (CI): 12.1–21.4] cases per 1000 person-years. Incidence rates				
37 38	39	in men and women were 17.7 (95% CI: 11.6–25.9) and 15.0 (95% CI: 9.8–22.2) cases per 1000				
39 40 41	40	person-years, respectively. Multivariable Cox proportional hazard analysis determined older age				
42 43	41	and obesity in women, and prediabetes in men to be the independent risk factors for developing				
44 45	42	type 2 DM.				
46 47 48	43	Conclusions: The incidence rate of type 2 DM in overweight and obese UAE nationals is high.				
49 50	44	In addition to screening, current strategies should strongly emphasize lifestyle modifications to				
51 52	45	decrease HbA1c and BMI levels in this high-risk population.				
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#### **Article Summary** 47

#### Strengths and limitations of this study 48

- This long-term longitudinal cohort study determines the incidence rate and risk factors of 49
- type 2 diabetes mellitus in overweight and obese United Arab Emirates nationals. 50
- 51 Anthropometric and laboratory data were obtained instead of self-reported information for
- the classification of risk factors and identification of incident cases. 52
  - 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.
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**Keywords:** incidence; obesity; risk factors; type 2 diabetes mellitus; United Arab Emirates 4.02

#### **INTRODUCTION** 60

A rapid worldwide increase has been observed regarding the prevalence of type 2 diabetes 61 mellitus (DM) and obesity. The United Arab Emirates (UAE) is one of the epicenters of this 62 epidemic. In a 2007 study, the age-standardized rates for prediabetes and type 2 DM among 63 UAE nationals were 24.2% and 29%, respectively.[1] However, in a 2019 study of young 64 Emirati men, the age-adjusted prevalence of prediabetes was much higher at 41.3%.[2] By 65 contrast, the prevalence of type 2 DM in North America and Western European countries are 66 well below 12%.[3-6] Notably, the number of people with type 2 DM in the UAE is estimated to 67 68 reach over two million by 2040.[7]

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### **BMJ** Open

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Overweight, defined as body mass index (BMI) between 25 and 29.99 kg/m<sup>2</sup> and obesity, 69 defined as BMI  $\geq$  30 kg/m<sup>2</sup>,[8] are known risk factors of type 2 DM.[9] More recently, the age-70 adjusted prevalence of overweight and obesity among Emirati men between the ages of 18 and 71 30 years was noted to be 27% and 30%, respectively.[2] In contrast, the corresponding 72 prevalence in another study was 23% and 10% among young Emirati women.[10] Both these 73 74 studies determined increased BMI to be strongly associated with diabetes. Both obesity and type 2 DM are well known cardiometabolic risk factors for vascular 75 complications, such as coronary heart disease and peripheral vascular disease. The prevalence of 76 77 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 78 global death rates for cardiovascular disease—204 per 100,000 for women and 309 per 100,000 79 for men.[11] 80 The rise in the incidence of diabetes and its complications among the UAE population has 81 been in parallel to the economic surge, resulting in growing urbanization and changing lifestyle 82 habits in the country. Nevertheless, over the last decade, UAE has diverted its attention and 83 endeavored to curb the burden of overweight and obesity, as well as to decrease the incidence of 84 85 associated conditions like diabetes. The Wegava program, a population-wide CVD screening program, was initiated in 2008[12] to enable individuals and healthcare providers to implement 86 87 CVD preventative measures. For continued assessment of the effectiveness of these local 88 interventions, ad hoc analysis of population data at pre-defined intervals over a long period is required. However, epidemiological studies on type 2 DM in the UAE have primarily focused on 89 its prevalence in the general population. [1, 2, 13] Despite having information from the Wegaya 90

program, data regarding the incidence of type 2 DM among obese and overweight UAE nationals

remains 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 2 DM and elucidate the effects of BMI on DM risk in adult UAE nationals
who are overweight and obese.

## 98 METHODS

## 99 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 April 1, 2008, and December 31, 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] The Research and Ethics Board of Tawam Hospital and the UAE University granted ethical approval for this study (CRD239/13). Informed consent was waived because patient records and information were anonymized and de-identified before analysis.

## 108 Subjects and procedures

109 The study population consisted of UAE nationals who were overweight and obese, aged between 110 18 and 75 years, who were seen at the outpatient departments at Tawam Hospital. The inclusion 111 criteria were patients without type 2 DM, defined as having glycated hemoglobin (HbA1c) 112 <6.5%; patients not taking antidiabetic medications; and those with BMI  $\ge$  25 kg/m<sup>2</sup> at baseline. 113 The exclusion criteria were patients with missing HbA1c data; those with a history of chronic 114 medical conditions, such as malignancy, human immunodeficiency virus infection, receiving Page 7 of 26

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steroids or immunosuppression medications, or stage 3–5 chronic kidney disease [estimated glomerular filtration rate (eGFR) of  $\leq$ 59 mL/min/1.73 m<sup>2</sup>].[16] Moreover, patients who had undergone 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 kg/m<sup>2</sup> (Fig. 1).

121 Definitions and measurements

Initial BMI was calculated from baseline height and weight measurements using the formula 122 123 weight (kg) divided by height squared  $(m^2)$ . Overweight and obesity were defined according to the World Health Organization classification [8] based on a BMI between 25 and 29.99 kg/m<sup>2</sup> 124 and  $\geq$ 30 kg/m<sup>2</sup>, respectively. Hypertension (HTN) was defined as systolic blood pressure  $\geq$ 140 125 mmHg, diastolic blood pressure  $\geq$ 90 mmHg, or the use of blood pressure medications.[17] The 126 definition of dyslipidemia was a serum total cholesterol (TC) of  $\geq 6.21$  mmol/L or if the patient 127 had a documented prescription of lipid-lowering medications.[18] Prediabetes was defined as 128 having an HbA1c level between 5.7% and 6.4%.[19] Current or prior history of smoking tobacco 129 was considered a positive smoking history. Patients with a documented history of coronary heart 130 disease, cerebrovascular disease, or peripheral vascular disease were identified to have a vascular 131 disease. 132

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## 134 Outcomes

Our study endpoint was incident type 2 DM, defined as either HbA1c ≥6.5% (based on the
definition of DM by the American Diabetes Association) or a diagnosis of type 2 DM established
by a physician at any time during the follow-up period.[19] The occurrence of incident type 2

DM was based on a review of all the electronic medical records until April 30, 2018. All laboratory analyses were performed at the Medical Laboratory Department of Tawam Hospital. An automated analyzer, Integra 400 Plus (Roche Diagnostics, Mannheim, Germany) was used to measure HbA1c levels. The HbA1c results were reported as a percentage using the International Federation of Clinical Chemistry-National Glycohemoglobin Standardization Program master equation.[20] Statistical analyses Using a formula by Rosner (2015),[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. Inter-sex differences were determined using the Mann-Whitney U test or the independent-samples *t*-test for non-normal and normal data, respectively. For categorical variables, Fisher's exact test (two-tailed) was used. Person-years 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% confidence interval (CI) was calculated per 1000 person-years by dividing the number of new cases of type 2 DM by the person-years at risk. Sex-specific incidence rates were calculated stratified by age and BMI categories. Kaplan-Meier time-to-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

160 predictors of incident type 2 DM in the entire cohort and stratified by sex. The following

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3 4	161	predictor variables were examined using univariable Cox modeling: age (categorized as <44
5 6	162	years and ≥44 years), sex, vascular disease, HTN, dyslipidemia, smoking, HbA1c (categorized as
7 8	163	<5.7% and $\geq$ 5.7%), and BMI (categorized as overweight and obese). All covariables with P <0.1
9 10 11	164	were included in the final multivariable model. A stepwise method based on likelihood ratios,
12 13	165	with entry and removal probabilities set at 0.05 and 0.10, respectively, was used to check the
14 15	166	results. The log-log plot was used to evaluate the proportional hazards assumption and was not
16 17 10	167	significant. A P value <0.05 was considered statistically significant. SPSS version 25 (IBM
18 19 20	168	Corp., Armonk, NY) was used to analyze the data.
21 22	169	
23 24	170	Patient and public involvement
25 26	171	There was no patient or public involvement in the design and conduct of the study
27 28	172	
29 30	172	
31 32	173	RESULTS
33 34	174	Characteristics of study participants
35 36	175	The baseline patient characteristics of the 362 subjects, are presented in Table 1. The ratio of
37 38	176	men to women was almost 1:1. At baseline, the mean age was $44.4 \pm 14.5$ years and the BMI
39 40 41	177	was 31.7 kg/m <sup>2</sup> [interquartile range (IQR) 27.6, 34.2 kg/m <sup>2</sup> ]. When stratified by sex, women
42 43	178	were older, had higher BMI and TC values, and had a higher prevalence of dyslipidemia at
44 45	179	baseline. Conversely, men had significantly higher baseline systolic and diastolic blood pressure
46 47	180	values, as well as a higher prevalence of smoking history.
48 49 50	181	
50 51 52	182	Table 1. Baseline characteristics of the study participants
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	Total (N = 362)	Men $(N = 170)$	Women (N =	
	$\frac{10 \tan (10 - 302)}{10 \tan (10 - 302)}$	M = 170	192)	P val
	Mean $\pm$ SD	Mean $\pm$ SD	Mean $\pm$ SD	
Women (%)	53.0	-	-	-
Age (years)	$44.4 \pm 14.5$	$40.2 \pm 14.7$	48.1 ± 13.2	<0.0
Anthropometric values				
BMI (kg/m <sup>2</sup> )	31.7 (27.6, 34.2)	29.6 (27.3, 32.8)	31.2 (27.9, 35.8)	0.0
SBP (mmHg)	$127.9 \pm 17.3$	$129.9 \pm 17.9$	$126.2 \pm 16.7$	0.04
DBP (mmHg)	✓ 76.9 ± 11.0	79.0 ± 11.3	75.1 ± 10.4	0.0
Laboratory values	6			
HbA1c (%)	$5.6 \pm 0.5$	5.6 ± 0.5	$5.6 \pm 0.5$	0.32
eGFR (mL/min/1.73 m <sup>2</sup> )	105.9 ± 16.8 (N = 359)†	$104.2 \pm 18.5 (N = 167)$ †	$107.4 \pm 15.1$	0.0
TC (mmol/L)	5.1 ± 0.9	4.9 ± 1.0	$5.2 \pm 1.0$	0.0
History of n (%)	0			
HTN	181 (50.0)	91 (53.5)	90 (46.9)	0.2
Dyslipidemia	141 (39.0)	52 (30.6)	89 (46.4)	0.0
Vascular disease	18 (5.0)	10 (5.9)	8 (4.2)	0.4
Smoking	56 (15.5)	54 (31.8)	2 (1.0)	<0.0

<sup>184</sup> systolic blood pressure, *DBP* diastolic blood pressure, *SD* standard deviation, *HbA1c* glycated

*†*The difference in N is because of the missing data.

187 \*Independent-samples t-test was used to calculate *P* values for continuous variables and Fisher's

- 188 exact test (two-tailed) for categorical variables.

190 Incidence

<sup>185</sup> hemoglobin, *TC* total cholesterol.

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3 4	191	After a median	follow-up time of 8.7 years	(IQR, 8.2-9.0 years), 47 in	cident cases of type 2 DM
5 6	192	were identified.	The overall incidence rate	of type 2 DM was 16.3 (95%	% CI: 12.1–21.4) cases per
7 8	193	1000 person-ye	ars, and the crude incidence	e over the same period was 1	13.0% (95% CI:
9 10 11	194	9.8–16.8%). Th	e incidence rate of type 2 I	DM increased with patient ag	ge (Fig. 2). When stratified
12 13	195	by sex, the incid	dence rates, were 17.7 (95%	6 CI: 11.6–25.9) and 15.0 (9	5% CI: 9.8–22.2) cases
14 15	196	per 1000 persor	-years in men and women,	respectively.	
16 17 19	197	Sex-specific	incidence rates, as stratified	d by BMI, were higher in ob	bese women (21.1, 95%
19 20	198	CI: 13.3–32.0 c	ases per 1000 person-years	) compared with overweight	t women (5.1, 95% CI:
21 22	199	1.3-14.0 cases	per 1000 person-years) (P =	= 0.013). Incidence rates for	type 2 DM were higher in
23 24 25	200	older women (2	1.3, 95% CI: 13.5–31.9 cas	es per 1000 person-years) c	ompared with younger
25 26 27	201	women (3.7, 95	% CI: 0.6–12.2 cases per 1	000 person-years) ( $P = 0.00$	8), (Fig. 3). However,
28 29	202	these difference	s were not significant amor	ng men.	
30 31	203	Analyses of ris	k factors		
32 33 34	204	We used multiv	ariable Cox regression anal	yses to examine the associa	tion of age, sex, vascular
35 36	205	disease, dyslipi	demia, smoking, HTN, BM	I, and HbA1c levels with the	e risk of developing type 2
37 38	206	DM in the entir	e study cohort, as well as b	etween men and women. Wi	hen adjusted for all other
39 40 41	207	variables, we ol	oserved that in men an HbA	1c level ≥5.7% [hazard ration]	o (HR) = 3.02, 95% CI:
41 42 43	208	1.20-7.62] and	in women age ≥44 years (H	IR = 4.84, 95% CI: 1.13–20	.73) and obesity (HR =
44 45	209	3.42, 95% CI: 1	.01–11.57) were significan	t risk factors for the develop	oment of type 2 DM (Table
46 47	210	2).			
48 49 50	211				
50 51 52	212	Table 2. Predic	tors of incident type 2 diab	etes mellitus in the study po	pulation
53 54		Predictor	Total (N = 362)	Men (N = 170)	Women (N = 192)
55 56 57					
58 59					10
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variables	HR (95% CI)	Р	HR (95% CI)	Р	HR (95%	Р
		value		value	CI)	value
Age						
<44 years	Reference		Reference		Reference	
$\geq$ 44 years	1.20 (0.59–2.44)	0.619	0.46 (0.17– 1.13)	0.135	4.84 (1.13– 20.73)	0.034
Sex						
Women	Reference					
Men	1.60 (0.83-3.05)	0.160				
History of HTN						
No	Reference		Reference		Reference	
Yes	1.78 (0.94–3.37)	0.075	1.89 (0.74– 4.81)	0.183	1.61 (0.67– 3.88)	0.292
History of dyslipidemia						
No	Reference		Reference		Reference	
Yes	1.32 (0.70–2.51)	0.393	2.01 (0.80– 5.04)	0.135	0.80 (0.34–1.90)	0.608
History of smoking						
No	Reference		Reference		Reference	
Yes	0.58 (0.23–1.49)	0.259	0.36 (0.12– 1.04)	0.060	4.88 (0.44– 54.08)	0.196
History of vascular disease			4.			
No	Reference		Reference		Reference	
Yes	1.57 (0.54–4.54)	0.408	2.46 (0.65–9.27)	0.183	0.60 (0.06–5.50)	0.647
HbA1c						
<5.7%	Reference		Reference		Reference	
≥5.7%	2.29 (1.16-4.52)	0.016	3.02 (1.20– 7.62)	0.019	2.30 (0.83– 6.39)	0.109
BMI						
Overweight	Reference		Reference		Reference	
Obese	1 86 (0 99_3 50)	0.053	1 17 (0 51_	0.713	342(101 -	0.040

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The results of this study revealed that the crude incidence of type 2 DM among the UAE 217 nationals who are overweight and obese was 1.49% per year during the 9-year study period. The 218 overall incidence rate was 16.3 per 1000 person-years. The independent predictors for the 219 development of type 2 DM were age (≥44 years) and obesity in women and prediabetes in men. 220 **Incidence** rates 221 222 The incidence rates of type 2 DM in high-risk populations differ worldwide. This difference is probably because of variations in insulin sensitivity related to the interaction between genetic 223 and environmental factors among different ethnic and population groups globally.[23] In a 224 225 United States study of predominantly overweight and obese African-American women, reported the incidence rate of type 2 DM was 15.2 per 1000 person-years [24] and is comparable to our 226 study. There has been only one other study that has assessed the incidence rate of type 2 DM in 227 the UAE.[13] The study was conducted over 12 months in Ajman, one of the emirates in the 228 UAE and revealed that the incidence rate for type 2 DM was 4.8 per 1000 person-years. 229 However, this study cannot be directly compared with our study because it was conducted on the 230 general population, and our study cohort comprised only overweight and obese UAE nationals. 231 **Risk factors** 232 This study revealed some interesting findings regarding the risk factors for developing type 2 233 DM in the high-risk UAE population. Sex-specific incidence rates of type 2 DM exhibited a 234 235 female preponderance, particularly older women. Nevertheless, these results were not unique and 236 were also observed among the general population of the UAE.[13] The higher incidence of type 2 DM among older women may be attributable to the high prevalence of overweight and obesity, 237 sedentary lifestyle, nutritional habits, and early menopause witnessed in our population.[25-28] 238

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Furthermore, individuals with high-normal HbA1c levels have a greater risk of developing 239 type 2 DM.[29] However, when we stratified our cohort by sex, the independent predictive 240 property of the HbA1c level was only observed in men. An explanation for this finding could be 241 the high prevalence of smoking history in men, as smoking has been shown to increase HbA1c 242 levels in individuals without diabetes.[30] Moreover, obesity in women was noted to be an 243 244 independent risk predictor for type 2 DM. Studies have demonstrated that the risk of developing type 2 DM is significantly higher in individuals with BMI  $\geq$  30 kg/m<sup>2</sup> compared with those with 245 normal BMI.[9] Therefore, interventions that promote weight loss and reduce BMI can reduce 246 247 the risk of developing type 2 DM. Notably, an eight-week exercise intervention program in a UAE student population demonstrated a significant reduction in BMI.[31] In addition, other 248 controlled intervention studies in high-risk populations[32,33] demonstrated a significant 249 250 decrease in the risk of type 2 DM consequent to BMI reduction.

# 1 251 Clinical and public health implications

In the UAE, much of the increase in type 2 DM has occurred because of social and economic 252 changes-from traditional, semi-nomadic lifestyle to sedentary, urban, and high-income 253 lifestyle. Consequently, the progression toward type 2 DM and cardiovascular disease starts early 254 in the UAE because of the high prevalence of childhood obesity.[1, 34] A 2015 study of Emirati 255 children and adolescents with a BMI  $\geq 25 \text{ kg/m}^2$  determined the prevalence of prediabetes and 256 type 2 DM to be 5.4% and 0.87%, respectively.[35] Therefore, reducing the burden of type 2 257 258 DM 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. 259

Furthermore, the results of this study highlight the need to focus on Emirati women as aspecific target group for obesity prevention programs. These programs should begin in childhood

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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 2 DM, wherein
pediatricians, primary care providers, dieticians, and specialists work collaboratively toward
promoting weight reduction and maintaining a healthy BMI in our population.

266 Strengths and limitations

The strengths of this study include the long duration of follow-up for our representative sample 267 of overweight and obese individuals in the second-largest city in the Abu Dhabi Emirate of UAE. 268 We used the anthropometric and laboratory data documented in patient charts instead of self-269 270 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 271 incident cases of type 2 DM instead of fasting plasma glucose that has more variability.[37,38] 272 Nonetheless, this study had some limitations. Baseline data regarding family history of type 2 273 DM, history of gestational diabetes, physical activity, and abdominal obesity were unavailable-274 all of which are well-researched risk factors for the development of DM. Nevertheless, a 275 prospective study can provide more robust results. Furthermore, data on additional confounders, 276 such as family history, changes in weight and physical activity, were unavailable for review. 277 278 Finally, 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. 279

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## 281 CONCLUSIONS

The overall incidence rate of type 2 DM in UAE nationals who are overweight and obese was
16.3 per 1000 person-years. Notably, prediabetes 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 emphasize BMI reduction through lifestyle modifications inthese high-risk groups.

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- 290 AUTHOR CONTRIBUTIONS
- 291 D.R. conceived the study and its design, performed literature search and drafted the initial

292 manuscript. S.A.S. was involved in data management, statistical analysis, interpretation of the

data, and critical review of the manuscript. R.D.G. assisted in statistical analysis, contributed to

the discussion, reviewed and edited the manuscript. J.A.K. performed a critical review of the

295 manuscript was involved in data interpretation and also guided the research methodology. All

authors read and approved the final manuscript.

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300 decision to publish, or preparation of the manuscript.

- <sup>301</sup> COMPETING INTERESTS
- 302 The authors declare that they have no competing interests.

# **303 PATIENT CONSENT FOR PUBLICATION**

304 Not required

# <sup>9</sup> 305 AVAILABILITY OF DATA AND MATERIALS

 $\frac{3}{22}$  306 The dataset used and analyzed during the current study is available from the corresponding

307 author on reasonable request.

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414	FIGURE LEGENDS
415	Figure 1. Flow chart for the inclusion and exclusion of patients
416	HbA1c glycated hemoglobin, eGFR estimated glomerular filtration rate, HIV human
417	immunodeficiency virus.
418	Figure 2. Overall incidence rate of type 2 diabetes mellitus (cases/1000 person-years) by
419	increasing age
420	Figure 3. Sex-specific incidence rate of type 2 diabetes mellitus (cases/1000 person-years)
421	stratified by BMI categories (A) and by age (B)
422	<i>BMI</i> body mass index.











Section/Topic	ltem #	Recommendation	Reported on page #			
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract	1			
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	2			
Introduction						
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	3, 4			
Objectives	3	State specific objectives, including any prespecified hypotheses	4, 5			
Methods						
Study design	4	Present key elements of study design early in the paper	5			
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	5, 6			
Participants	6	(a) Give the eligibility criteria, and the sources and methods of selection of participants. Describe methods of follow-up	6, 7			
		(b) For matched studies, give matching criteria and number of exposed and unexposed	NA			
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	6, 7			
Data sources/ measurement	8*	For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group	6,7			
Bias	9	Describe any efforts to address potential sources of bias	NA			
Study size	10	Explain how the study size was arrived at	7			
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why	7			
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	7			
		(b) Describe any methods used to examine subgroups and interactions	7			
		(c) Explain how missing data were addressed	5			
		(d) If applicable, explain how loss to follow-up was addressed	5			
		(e) Describe any sensitivity analyses	NA			
Results						

# STROBE 2007 (v4) Statement—Checklist of items that should be included in reports of cohort studies

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Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility, confirmed	5, 6
		eligible, included in the study, completing follow-up, and analysed	
		(b) Give reasons for non-participation at each stage	5
		(c) Consider use of a flow diagram	Figure 1
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential	Table 1
		confounders	
		(b) Indicate number of participants with missing data for each variable of interest	Table 1
		(c) Summarise follow-up time (eg, average and total amount)	10
Outcome data	15*	Report numbers of outcome events or summary measures over time	10
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence	Table 2
		interval). Make clear which confounders were adjusted for and why they were included	
		(b) Report category boundaries when continuous variables were categorized	Table 2
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	NA
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	10
Discussion			
Key results	18	Summarise key results with reference to study objectives	12
Limitations			
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from	12–14
		similar studies, and other relevant evidence	
Generalisability	21	Discuss the generalisability (external validity) of the study results	14
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on	15
		which the present article is based	

\*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

**Note:** An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.