







# Incidence of Insulin-requiring Diabetes in the US Military

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# SHORT COMMUNICATION

# Incidence of insulin-requiring diabetes in the US military

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#### Abstract

Aims/hypothesis The aim of the study was to determine age- and race-related, and overall incidence rates of insulin-requiring diabetes in adults in the US military.

Methods Electronic records for admissions to US military and Tricare hospitals during 1990–2005 and visits to military clinics during 2000–2005 were identified using the Career History Archival Medical and Personnel System at the Naval Health Research Center, San Diego, CA, USA. Population data were obtained from the Defense Manpower Data Center and Defense Medical Epidemiology Database. Results In men there were 2,918 new cases of insulinrequiring diabetes in 20,427,038 person-years at ages 18–44 years (median age 28 years) for a total age-adjusted incidence rate of 17.5 per 100,000 person-years (95% CI 16.4–18.6). Incidence rates were twice as high in black men as in white men (31.5 vs 14.5 per 100,000, p<0.001). In

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C. Ricordi Diabetes Research Institute, University of Miami, Miller School of Medicine, Miami, FL, USA women there were 414 new cases in 3,285,000 personyears at ages 18–44 years (median age 27 years), for a total age-adjusted incidence rate of 13.6 per 100,000 (95% CI 12.4–14.9). Incidence rates were twice as high in black women as in white women (21.8 vs 9.7 per 100,000, p<0.001). In a regression model, incidence of insulinrequiring diabetes peaked annually in the winter–spring season (OR 1.46, p<0.01). Race and seasonal differences persisted in the multivariate analysis.

Conclusions/interpretation Differences in incidence rates by race and season suggest a need for further research into possible reasons, including the possibility of a contribution from vitamin D deficiency. Cohort studies using prediagnostic serum 25-hydroxyvitamin D should be conducted to further evaluate this relationship.

**Keywords** Incidence rates · Insulin-requiring diabetes · Race · Seasonality · Sex · Vitamin D · Young adults

# Abbreviations

CHAMPS Career History Archival Medical and

Personnel System

DMED Defense Medical Epidemiology Database

## Introduction

Insulin-requiring diabetes is the second most common chronic disease of childhood in the USA after asthma [1]. There are thought to be 15,000–30,000 new cases of insulin-requiring diabetes in childhood in the USA annually. However, there are few reports on population-based incidence data for the disease [1–4]. This limits the validity of current estimates of the burden of disease. The present study reports the age-



specific incidence of insulin-requiring diabetes in adult men and women, and its association with race and season.

#### Methods

Persons with insulin-requiring diabetes are not allowed to join the US military. As a result, first hospitalisations or clinic visits during military service represent incident cases. Hospitalisation data from military hospitals were obtained from the Tricare Executive Information and Decision Support System, Standard Inpatient Data Record (SIDR), which includes discharge summaries of all admissions. Hospitalisation data from civilian Tricare hospitals were obtained from the Tricare Health Care Service Record (HCSR) database. Outpatient records of treatment in military clinics were obtained from the Standard Ambulatory Data Record (SADR) database, which included outpatient visits to all military clinics worldwide in 2000–2005. Tricare hospitals and military outpatient clinics have participated in the current records system since 2000.

Electronic records of inpatient encounters were identified for service members aged 18–44 years who were hospitalised with a discharge diagnosis of insulin-requiring diabetes in any military hospital during 1990–2005, in a civilian Tricare hospital during 2000–2005, or treated in any outpatient military clinic worldwide during 2000–2005.

Data on hospitalisations and clinic visits were entered on a monthly basis into the Career History Archival Medical and Personnel System (CHAMPS) maintained by the Naval Health Research Center (San Diego, CA, USA). A CHAMPS history consists of an individual record for each service member. It includes medical and career events and captures admissions to any military or Tricare hospital, or visits to any military outpatient clinic in the world.

Case definition A case was defined as: (1) any individual who had a hospital discharge diagnosis of insulin-requiring diabetes (ICD9-CM Codes 250.X1 or 250.X3, where X is 0-9; www.icd9data.com/2007/Volume1) in either a US Department of Defense medical treatment facility or a Tricare civilian hospital; (2) any individual who had at least three outpatient visits listing type 1 diabetes as the diagnosis during any 12 month period; or (3) any individual who had at least seven outpatient visits with a diagnosis of insulin-requiring diabetes during their military career. The reason for requiring multiple clinic visits for identification as a case is that insulinrequiring diabetes was occasionally recorded as a tentative diagnosis explaining a clinic visit pending confirmation by laboratory data. A review of records revealed that when there were three or more clinic visits for insulin-requiring diabetes in a 12 month period, the individual generally had more than ten subsequent visits for evaluation and treatment of insulinrequiring diabetes or received a discharge from service due to

Denominator data were obtained from the Defense Medical Epidemiology Database (DMED; Silver Spring, MD, USA) of the US Army and the Defense Manpower Data Center-West (Seaside, CA, USA). All records were maintained in CHAMPS, which is a federally designated system of records, with strict precautions regarding maintenance of confidentiality of patient health information.

Incidence rates in the Department of Defense population were adjusted to the US standard population in 2000 [5] using the direct method [6]. The 95% CI were calculated using the Poisson distribution [6]. Multiple logistic regression was used to adjust for age, sex, race and season. The criterion for statistical significance of all analyses in the study was an alpha level of 0.05, two-tailed.

Multiple regression analysis was performed using all cases during 2000–2004 and a probability sample of non-cases who were on active duty in the military at the time of first diagnosis of the cases. All analyses were performed using SAS-PC, version 7.0 PROC FREQ and PROC LOGISTIC (SAS Institute, Cary, NC, USA).

#### Results

In men, there were 2,918 newly diagnosed cases of insulinrequiring diabetes at ages 18–44 years (median age 28 years) in 20,427,038 person-years at risk (Table 1). The overall age-standardised incidence rate in men was 17.5 per 100,000 person-years. Incidence rates in black men were approximately double those in white men (p< 0.001), and rose consistently with age (Table 1).

In women, there were 414 newly diagnosed cases of insulin-requiring diabetes at ages 18–44 years (median age 27 years) in 3,285,000 person-years at risk (Table 1), corresponding to an age-standardised incidence rate of 13.6 per 100,000 person-years. Incidence rates in black women were approximately double those in white women (p< 0.001, Table 1). Incidence rates in men and women of other ethnicities were generally intermediate between those of white and black people (Table 1). The higher rates in black persons and in late winter (see Electronic Supplementary Material [ESM] Fig. 1) persisted after adjustment for age and sex, indicating that being black and winter season were independently associated with risk of insulin-requiring diabetes (Table 2).

#### Discussion

This study found substantially higher incidence rates of insulin-requiring diabetes in blacks and marked seasonal



**Table 1** Annual age-specific and age-adjusted incidence rates of insulin-requiring diabetes per 100,000 person-years by race and sex (active duty service members) (Department of Defense, 1990–2005)

Age (years)	White			Black			Other		
	No. of new cases	No. of person-years at risk	Annual rate per100,000 person-years	No. of new cases	No. of person-years at risk	Annual rate per 100,000 person-years	No. of new cases	No. of person-years at risk	Annual rate per 100,000 person-years
Men									
<20	122	1,203,505	10.1	26	269,053	9.7	3	130,811	2.3
20–24	537	4,979,268	10.8	173	1,166,034	14.8	32	544,029	5.9
25-29	363	3,212,950	11.3	152	860,290	17.7	21	356,393	5.9
30-34	321	2,465,765	13.0	197	707,407	27.8	23	263,237	8.7
35–39	334	2,058,562	16.2	241	584,540	41.2	31	216,669	14.3
40-44	195	1,045,387	18.7	128	241,992	52.9	19	121,146	15.7
Total	1,872	14,965,437	12.5	917	3,829,316	23.9	129	1,632,285	7.9
Age-adjusted rate			14.5			31.5			13.5
95% CI			13.6-15.4			29.6-33.5			11.6-15.7
Women									
<20	11	193,854	5.7	20	89,624	22.3	4	31,053	12.9
20–24	72	711,972	10.1	41	347,665	11.8	5	113,790	4.4
25–29	47	435,330	10.8	43	247,970	17.3	10	64,861	15.4
30–34	32	288,039	11.1	56	180,873	31.0	7	36,145	19.4
35–39	27	222,473	12.1	19	130,233	14.6	2	22,807	8.8
40-44	8	107,953	7.4	10	50,031	20.0	0	10,327	0.0
Total	197	1,959,621	10.1	189	1,046,396	18.1	28	278,983	10.0
Age-adjusted rate			9.7			21.8			11.1
95% CI			8.4-11.2			18.9-25.1			7.4-16.0

The total annual age-adjusted incidence rate for the whole population was 17.5 (95% CI 16.4–18.8) per 100,000 person-years in men and 13.6 (95% CI 12.4–14.9) per 100,000 person-years in women. The p values for the difference in incidence rates between white and black men was <0.001, and in women it was also <0.001. The median age of the cases was 28 years in men and 27 years in women

**Table 2** Estimated relative risk of insulin-requiring diabetes according to age, sex, race, and quarter (Department of Defense service members, 2000–2004)

Covariate	R	SE	t	OR	95% CI	p value
Age (years)						
18-24				1.00	Reference	
25–34	0.129	0.067	1.9	1.14	1.00-1.30	0.05
≥35	1.657	0.072	23.0	5.25	4.56-6.04	< 0.001
Sex						
Male				1.00	Reference	
Female	-0.124	0.083	-1.5	0.88	0.75 - 1.04	0.14
Race						
White				1.00	Reference	
Black	0.748	0.060	12.5	2.11	1.88-2.38	< 0.01
Other	0.261	0.120	2.2	1.30	1.03-1.64	0.03
Quarter						
July-September				1.00	Reference	
October-December	0.055	0.084	0.7	1.06	0.90-1.25	0.52
January-March	0.378	0.078	4.8	1.46	1.25 - 1.70	< 0.01
April–June	0.213	0.081	2.6	1.24	1.06-1.45	< 0.01



variation in incidence in all races combined, which peaked in late winter through early spring and were lowest in summer and early autumn (Table 2). There are few previous reports on racial differences in insulin-requiring diabetes incidence rates in young adults. A study of Navy-enlisted men during 1974–1988 found a higher incidence rate in blacks than whites (28.4 vs 20.2 per 100,000 person-years, respectively; p<0.05) [2]. Data presented here indicate that insulin-requiring diabetes occurs beyond childhood at a substantial incidence rate and that it appears to be more common in black than in white people at ages 18–44 years.

It is possible that some of the individuals in the population identified as cases of insulin-requiring diabetes may actually have type 2 diabetes. In addition, the observed age-related increase in incidence of insulin-requiring diabetes (Table 1) suggests that some cases of type 2 diabetes may have been included in the analysis because they were being given insulin. Therefore, caution should be exercised in drawing conclusions about whether insulin-requiring diabetes in this population is specifically type 1 or type 2. On the other hand, most cases were hospitalised and the diagnosis of all was type 1 diabetes, indicating that all cases were receiving insulin at the time of hospital discharge.

Previous research has demonstrated an association between BMI and risk of insulin-dependent diabetes [7]. However, data on BMI were not available in the database used for the present study or any database available to the investigators for this population. Caution should be exercised when interpreting the results, as the higher risk observed in later age and in blacks may be due to higher BMI in these groups. Furthermore, the military population is not a probability sample of the US population, so the results cannot be generalised to the US population without caution.

Aetiology Several theories have been put forward in an effort to explain the epidemiology and cause of insulinrequiring diabetes, yet the identification of a causal factor has been elusive. The armed services actively promotes exercise, physical fitness and weight control, and as a result, might be expected to have a lower than average incidence rate of type 2 diabetes. However, there is less evidence that physical activity or weight influences the incidence of type 1 diabetes requiring insulin in children [8], unlike diabetes not requiring insulin. Differences in body mass between white and black populations have been proposed as possibly accounting for the higher incidence rate of type 2 diabetes in black people [9]; however, this explanation is not wholly applicable to young insulinrequiring diabetes patients, who are usually not overweight [8]. Some evidence suggests that blacks diagnosed with insulin-requiring diabetes are less likely than whites to

have antibodies against islet-cell antigens, and that they may be more likely than whites to have type 2 diabetes presenting acutely as insulin-requiring diabetes [10].

Winter season and black skin pigmentation substantially reduce the photosynthesis of vitamin D<sub>3</sub> in the skin, creating a high prevalence of vitamin D deficiency. The markedly higher incidence of insulin-requiring diabetes in black than in white people in the present study suggests the possibility that vitamin D deficiency might play some role in the differential incidence of insulin-requiring diabetes according to race, as suggested by previous research [11, 12]. If vitamin D deficiency is a contributor to insulin-requiring diabetes [11, 12], it could help explain the seasonal and racial differences in incidence rates.

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**Duality of interest** The authors declare that there is no duality of interest associated with this manuscript.

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## 13. SUPPLEMENTARY NOTES

# 14. ABSTRACT

Type 1 diabetes mellitus (T1DM) is a disease of concern in young adults and children. Further research would be desirable to define the role of race and seasonality in incidence of T1DM in young adults. Seasonality and racial disparities in incidence suggest that T1DM may be associated vitamin D inadequacy. If so, public health action to reduce its incidence may be feasible with supplementation and vitamin D fortification of foods.

Seasonal patterns and racial disparities in diabetes incidence that exist in military populations with uniform access to medical care may indicate a role of vitamin D status in diabetes control. A protective association between vitamin D adequacy and risk of T1DM has implications for active-duty military personnel at risk for this condition.

# 15. SUBJECT TERMS

vitamin D, childhood type 1 diabetes

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