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## Regional Differences in the Management of Cardiovascular Risk Factors among Adults with Diabetes: An Evaluation of the Diabetes Collaborative Registry

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### Abstract

**Aims:** To compare cardiovascular risk factor control in adults with diabetes participating in a national diabetes registry to those in the general population and to ascertain regional differences in diabetes care.

**Methods:** Adults with diagnosed diabetes in the Diabetes Collaborative Registry (DCR) were compared with those in the National Health and Nutrition Examination Survey (NHANES) from 2015–2016; standardized mean difference (SMD) $>0.2$  defined significance. Regional differences were assessed in the DCR cohort;  $p<.05$  defined significance.

**Results:** The DCR cohort was older (61 vs. 57 years, SMD=0.38), more insured (99.7% vs. 91.0%, SMD=0.42), and less ethnically diverse (83% non-Hispanic white vs. 76%, SMD=0.30) compared with NHANES. The proportion of overweight/obesity, A1c $<7\%$  ( $<53$  mmol/mol), and BP $<140/90$  were similar, but DCR participants had higher proportion with LDL $<2.59$  mmol/L (61% vs. 41%, SMD=0.39) and fewer tobacco users (17% vs. 32%, SMD=0.35). Regionally, obesity, lack of glycaemic control, and tobacco use were highest in the Midwest, BP control was the lowest in the South, and LDL control was lowest in the Northeast.

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#### Author contributions

L.J.T. and S.N.M. interpreted data and drafted the manuscript. Y.T. performed the statistical analysis. All remaining authors contributed to study concept and design, data interpretation, and critically revised the manuscript. S.N.M. is guarantor of this work and as such, takes responsibility for its contents.

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**Conclusions:** Significant regional differences in diabetes care delivery and outcomes were identified using a national diabetes registry. Serial analyses of the DCR may supplement national evaluations to deepen our understanding of diabetes care in the US.

## Keywords

Diabetes Mellitus; Cardiovascular Disease; Risk Factors; Pharmacoepidemiology; Population Health; Health Services Research

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## 1. Introduction

Diabetes mellitus remains a major cause of morbidity and premature mortality in the United States (US).<sup>1</sup> Failure to adequately manage cardiovascular risk factors contributes significantly to negative outcomes, including heart disease and stroke.<sup>2–4</sup> National registries and surveys have been developed to identify opportunities for improving cardiovascular disease risk factor management in persons with diabetes.

The National Health and Nutrition Examination Survey (NHANES) has evaluated nationally-representative US cohorts for 20 years, informing national trends in the management and outcomes of persons living with diabetes.<sup>5, 6</sup> However, the biennial assessments were not designed to inform geographic differences in diabetes care which could inform health policy at regional levels.<sup>2, 7</sup> Analyses of health system data, insurance claims information, and state and local registries have evaluated diabetes care in specific catchments, but variable study designs prevent evaluation of broader regional differences in the US population.<sup>6–11</sup>

The Diabetes Collaborative Registry (DCR) is the largest, multi-site diabetes registry in the US, collecting electronic health data of persons with diabetes from outpatient visits to primary care, endocrinology, and cardiology practices.<sup>12</sup> This registry may inform regional differences in the management of cardiovascular disease risk in adults with diabetes to supplement broader, national evaluations conducted on a biennial basis. However, the extent to which DCR participants are representative of the general population of persons with diabetes in the US is not known.

We compared persons with diagnosed diabetes participating in the DCR to the general population as assessed by NHANES using 2015–2016 data. Then, we evaluated geographic differences in demographic and clinical characteristics of the DCR cohort according to US Census region.

## 2. Methods

### 2.1. Data Sources

**Diabetes Collaborative Registry (DCR)**—The DCR is a prospective, outpatient, quality improvement registry of persons with diabetes and prediabetes, seen in ambulatory care settings across the US. With administrative oversight by the American College of Cardiology’s National Cardiovascular Data Registries, DCR was launched in 2014 as a

collaborative effort by several partner professional societies and includes primary care, endocrinology, and cardiology practices.

Individuals in the DCR have physician-diagnosed diabetes based on standardized criteria used in clinical care. Longitudinal patient data are collected at the point of care through an automated system integration solution that periodically extracts relevant data elements from electronic health records (EHRs), including demographics, comorbidities, clinical factors, laboratory values, and medications. Data collection is standardized using established definitions, uniform data entry and transmission, and quality checks. As a specialty registry recognized by the Center for Medicare and Medicaid Services (CMS), the DCR is periodically audited to ensure the accuracy of data for participation in federal incentive programs. For persons with multiple clinic visits between 2015–2106, data from the most recent visit were analysed. DCR participation requires no data collection beyond that of routine clinical care, and a waiver of written informed consent was granted by Chesapeake Research Review.

**NHANES**—NHANES is a program of studies designed to assess the health and nutrition of adults and children in the US. NHANES uses stratified, multistage, probability-cluster techniques to ensure a representative sample of the nation’s non-institutionalized civilian population. NHANES data include interview data for demographic, socioeconomic, and health-related questions in addition to examination data for physiological and laboratory measurements conducted by trained medical personnel. All participants provided written informed consent and the research ethics board of the National Center for Health Statistics approved all NHANES protocols.

## 2.2. Study Population

Adults in both cohorts aged 18–75 years with a diabetes diagnosis were included; those with a diagnosis of gestational or prediabetes were excluded. Only participants with a measure of glycated haemoglobin (A1c) were included in the analysis.

For DCR, participants with a diagnosis code from the International Classification of Diseases, 9th Revision (ICD-9) or 10th Revision (ICD-10) indicating type 2 diabetes or type 1 diabetes were included. Similarly, relevant ICD codes from each participant’s problem list were used to identify physician-diagnosed hypertension and dyslipidaemia. To better represent the general population with diabetes, DCR practices were restricted to primary care settings based on CMS’s Healthcare Provider Taxonomy Codes for internal medicine, family practice, and general practice, resulting in inclusion of 212 sites. The DCR cohort was divided according to US Census Region (Northeast, Midwest, South, and West) based on location of the clinical practice.

For NHANES, a ‘yes’ response to the question ‘Other than during pregnancy, {have you/has Sample Person}/{Have you/Has Sample Person} ever been told by a doctor or health professional that {you have/{he/she/Sample Person} has} diabetes or sugar diabetes?’ identified participants with a diabetes diagnosis. NHANES participants who had laboratory evidence of diabetes but did not report a diabetes diagnosis were classified as “undiagnosed diabetes” and excluded from the analysis. Similarly, respondents were asked if they had ever

been told by a doctor or other health professional that they had hypertension or dyslipidaemia. A 'yes' response defined NHANES participants as being diagnosed with hypertension and/or dyslipidaemia, respectively.

### 2.3. Data Collection

Clinical data were collected by automated extraction from EHRs of clinical sites participating in DCR and through interview and laboratory testing in NHANES. Body mass index (BMI) categories were defined as underweight (BMI <18.5 kg/m<sup>2</sup>), normal weight (BMI 18.5–24.9 kg/m<sup>2</sup>), overweight (BMI 25.0–29.9 kg/m<sup>2</sup>) and obese (BMI ≥30 kg/m<sup>2</sup>). Blood pressure (BP) control was defined as systolic BP (SBP) <140 mmHg and diastolic BP (DBP) <90 mmHg, while lipid control was defined as LDL cholesterol (LDL) <2.59 mmol/L. Control of A1c, BP, and cholesterol ("ABC control") was defined as composite attainment of A1c <7.0% (<53 mmol/mol), BP <140/90 mmHg and LDL <2.59 mmol/L.

Use of glucose-lowering, antihypertensive, and lipid-lowering medications was analysed in relation to control of the respective CVD risk factor. Medication data were extracted directly from EHRs for DCR participants using the most recent observation. NHANES respondents were asked about medication use in the past 30 days from the time of the interview.

### 2.4. Statistical Analysis

Comparisons between DCR and NHANES are followed by regional analyses of the DCR cohort. NHANES analyses were weighted to account for the NHANES complex design. The weights were provided by NCHS as they are designed to account for unequal probabilities of selection and non-response. Demographics, clinical characteristics, and medication use were reported as percentages (categorical variables) or means±SD (continuous variables). *P* values were generated via Chi-square or *t* tests, as appropriate.

Due to the large difference in sample size between the DCR and NHANES cohorts, standardized mean difference (SMD) was calculated for each comparison by dividing the difference in means by the pooled standard deviation; these values are presented alongside or in lieu of *p* values. SMD was calculated for categorical variables using the multivariate Mahalanobis distance method as described by Yang and Dalton.<sup>13</sup> We characterized effect sizes based on SMD as large (SMD >0.8), moderate (SMD >0.5–0.8), and small (SMD >0.2–0.5).<sup>14</sup> For regional analyses of the DCR cohort, *p* values <.05 defined significance. Statistical analyses were performed using SAS version 9.4 (SAS Institute, Cary, NC).

## 3. Results

### 3.1. Participant characteristics

Figure 1 presents sample derivation for the DCR and NHANES cohorts. The DCR cohort included 67,433 adults with diabetes (98% type 2 diabetes) seen in a primary care setting between 2015–2016 with an available A1c value. In the DCR cohort, 18% of adults received diabetes care in the Northeast, 17% in the Midwest, 62% in the South, and 3% in the West. The weighted NHANES 2015–2016 sample represented 21,479,573 people in the US population with an available A1c value. Table 1 presents demographic and clinical

characteristics of the DCR and NHANES cohorts. The DCR cohort was older ( $61\pm 10$  years vs.  $57\pm 12$  years,  $SMD=0.38$ ) and comprised of a larger percentage of non-Hispanic white (83% vs. 76%,  $SMD=0.30$ ). Nearly all DCR participants had insurance compared with NHANES participants (99.7% vs. 91.0%,  $SMD=0.42$ ).

Regional differences within the DCR cohort are presented in Table 2. The West region comprised the oldest cohort and the Northeast comprised the youngest ( $p<.0001$ ). The percentage of men differed by region with the smallest percentage in the South region. A majority of DCR participants were non-Hispanic white across all regions; the largest percentage of non-Hispanic blacks were in the South ( $p<.0001$ ). Insurance coverage rates were high across all regions (99–100%).

### 3.2 Cardiovascular risk factor control

The average BMI was similar and in the obese range for both DCR and NHANES participants with diabetes ( $33.3\pm 7.3$  kg/m<sup>2</sup> vs.  $32.2\pm 7.6$  kg/m<sup>2</sup>,  $SMD=0.05$ ). The prevalence of overweight and obesity was 90.6% in the DCR cohort and 90.2% in the NHANES sample ( $SMD=0.03$ ). There were significant regional differences in weight status in the DCR cohort with the highest percentage of obesity in the Midwest (69.6%,  $p<.0001$ ).

Mean A1c was similar between DCR participants and NHANES participants ( $7.5\pm 2.2\%$  [ $58.1\pm 24.4$  mmol/mol] vs.  $7.4\pm 1.8\%$  [ $57.3\pm 20.0$  mmol/mol],  $SMD=0.03$ ). The distributions of A1c <7% (<53 mmol/mol), A1c 7–9% (53–75 mmol/mol), and A1c >9% (>75 mmol/mol) were similar between both cohorts ( $SMD=0.16$ ). In the DCR cohort, the mean A1c was higher in the Midwest compared with all other regions ( $p<.0001$ ).

DCR participants had a higher percentage of diagnosed hypertension compared with NHANES participants ( $SMD=0.37$ ). The percentage with BP<140/90 was similar between cohorts ( $SMD=0.09$ ). Regional differences in blood pressure control were present in the DCR cohort. Overall, the South region had the lowest percentage attaining BP<140/90 compared with all other regions ( $p<.0001$ ). DCR participants in the West had the highest percentage of diagnosed hypertension.

DCR participants were also more likely to have a diagnosis of dyslipidaemia compared with NHANES participants (83% vs. 63%,  $SMD=0.48$ ). There were significant differences in lipid parameters between the cohorts. Sixty-one percent of DCR participants compared with 41% of NHANES participants had LDL<2.59 mmol/L ( $SMD=0.39$ ). Significant regional differences in lipid control were present in the DCR cohort. Overall, the South and Northeast regions had the lowest percentage attaining LDL<2.59 mmol/L compared with all other regions, while DCR participants in the West were most likely to have a diagnosis of dyslipidaemia and attain LDL<2.59 mmol/L ( $p<.0001$  for all comparisons).

The DCR had a significantly higher percentage of adults with diabetes attaining the composite measure of A1c<7% (<53 mmol/mol), BP<140/90, and LDL<2.59 mmol/L (“ABC control”) compared with NHANES participants (18.8% vs. 6.0%,  $SMD=0.40$ ). Tobacco use was lower in the DCR compared with NHANES (18% vs. 32%,  $SMD=0.35$ ). Within the DCR, ABC control differed across regions: 14% in the Midwest, 19% in the

West, 20% in the South, and 21% in the Northeast ( $p<.0001$ ). Tobacco use was highest in the Midwest and lowest in the Northeast.

### 3.3. Treatment characteristics

Tables 1 and 2 describe medication use in the DCR and NHANES cohorts and by Census region for the DCR cohort, respectively. The use of insulin (18% in DCR vs. 27% in NHANES) and oral hypoglycaemic agents (56% in DCR vs. 71% in NHANES) differed between the cohorts (SMD=0.22–0.33). There were significant differences in diabetes management when stratified by A1c level (Figure 2). The use of diabetes medications was similar between DCR and NHANES for those with A1c>9% (>75 mmol/mol) (SMD=0.16). Within the DCR cohort, there were significant regional differences in the use of diabetes medications, with greatest use of insulin and oral hypoglycaemic agents in the West ( $p<.0001$ ). Regional differences in medication use according to metabolic control are available in the Supplemental Appendix (Supplemental Figure 1).

The use of anti-hypertensive medications was greater in NHANES participants compared with DCR participants (95% vs. 79%, SMD=0.50). DCR participants were less likely to be prescribed anti-hypertensive therapy compared to NHANES participants in the setting of either diagnosed hypertension (88% vs. 95%, SMD=0.27) or BP 140/90 (85% vs. 92%, SMD=0.24) (Supplemental Figure 2a). DCR participants in the West had the highest use of anti-hypertensive medications across all regions ( $p<.0001$ ). Similar regional patterns were found among those with BP 140/90 mmHg: West 98%, Midwest 93%, South 83%, and Northeast 80% ( $p<.0001$ ) (Supplemental Figure 2b).

The use of cholesterol-lowering medications was similar in the DCR and NHANES cohorts (SMD=0.08). DCR participants were less likely to be prescribed lipid lowering therapy compared with NHANES participants in the setting of either diagnosed dyslipidaemia (78% vs. 90%, SMD=0.33) or LDL 2.59 mmol/L (60% vs. 77%, SMD=0.38) (Supplemental Figure 3a). In the DCR cohort, adults with diabetes in the West had the highest use of cholesterol-lowering medications followed by adults in the Midwest, South and Northeast. Similar regional patterns were found among those with LDL 2.59 mmol/L: West 75%, Midwest 68%, South 60%, and Northeast 57% ( $p<.0001$ ) (Supplemental Figure 3b).

## 4. Discussion

We report significant regional differences in the prevalence, diagnosis, and management of cardiovascular risk factors among adults with diabetes participating in the largest outpatient diabetes registry in the US. While metabolic control was better in the DCR with 1 in 5 attaining composite “ABC control” compared with 1 in 16 in the general US population with diabetes, these data confirm the high prevalence of CVD risk factors in adults with diabetes. One in 5 adults with diagnosed diabetes and A1c>9% were not prescribed any glucose-lowering medication, and one in 4 adults with comorbid dyslipidaemia were not on a cholesterol-lowering medication. DCR participants were similar in terms of many demographic parameters to the general population with diabetes as assessed by NHANES. DCR participants were older and less ethnically diverse, and future analyses may consider weighting the DCR cohort to account for these differences.

Lack of health insurance is associated with undiagnosed health conditions, inadequate management, and poorer health outcomes.<sup>15</sup> The DCR cohort had nearly complete insurance coverage compared with the general population with diabetes whose rate of insurance coverage (~91%) was consistent with Census estimates. Cohorts like DCR, drawn from EHR data from ambulatory clinical care settings, are more likely to be insured as they represent those with greater access to medical care services.<sup>16, 17</sup> While the higher percentage of diagnosed hypertension and dyslipidaemia in the DCR cohort may suggest greater patient engagement with the healthcare system, medication use was equal, if not higher, in the general population with diabetes. There exists significant heterogeneity in insurance coverage for diabetes medications and medical devices amongst the myriad of commercial and public insurance plans complicated further by the use of various forms of supplemental or secondary insurance. The specific impact of insurance coverage on the receipt of recommended diabetes care and attainment of CVD risk factor control warrants further investigation in the DCR and NHANES cohorts.

Previous studies of the general population have demonstrated lower CVD risk for those living in the West and Northeast regions of the US compared to the South and Midwest regions and regional analyses of the DCR cohort confirm similar findings among those with diabetes.<sup>2, 4, 8, 18</sup> Adults with diabetes in the West were more likely to be diagnosed with hypertension and dyslipidaemia as well as be prescribed insulin, antihypertensive and lipid-lowering medications. While this may relate to the older age of the West cohort, it may also reflect greater attention to personal health or engagement with their primary care providers. Adults from the West were most likely to attain LDL<2.59 mmol/L, but adults from the Northeast were more likely to attain A1c<7% (<53 mmol/mol), BP<140/90, and composite ABC control. The Northeast region had relatively low levels of diagnosed dyslipidaemia with the lowest percentage attaining LDL<2.59 mmol/L, suggesting under-recognition of this important CVD risk factor in the Northeast. Future analyses are needed to confirm this finding, which has not been reported in the general population.

By contrast, adults with diabetes from the Midwest were noted to have the highest rates of obesity, A1c >9% (>75 mmol/mol), and tobacco use in addition to the lowest attainment of composite ABC control. These findings are particularly concerning given the high rates of medication use in the Midwest. Clinical inertia and medication non-adherence are commonly reported causes for discrepancies between medication prescriptions and poor metabolic control and warrant further investigation.<sup>19, 20</sup> Average A1c in the South was among the lowest in the US which mirrors findings by Egede et al, who reported that veterans with diabetes residing in the South had the lowest A1c values.<sup>11</sup> BP control was worst in the South which may relate to low rates of antihypertensive medication use, access to health care, clinical inertia, or genetic factors, noting the higher percentage of women and non-Hispanic blacks in this region. While future analyses may adjust for sex and race to account for possible genetic differences, efforts to improve antihypertensive use in the South are likely to be impactful. Lower rates of exercise, less healthful dietary behaviours, and persistently higher levels of tobacco use in the Midwest and South are well-described and likely contribute to the higher prevalence of CVD risk factors in these regions.<sup>21</sup>



The study has limitations that merit further discussion. First, diagnosed medical conditions were defined differently in the DCR and NHANES. Problem lists from EHRs, as used by DCR, have unique advantages and weaknesses compared with patient self-report, as used by NHANES, which may have influenced the findings.<sup>6, 7, 22–24</sup> Our analysis aimed to maximize the likelihood of a known diagnosis of diabetes, hypertension, or dyslipidemia by requiring specific ICD-10 codes for these conditions. We chose not to include medication use without an associated ICD diagnosis code as indicative of a diagnosis, as these medications may be used for prevention of disease (e.g. prediabetes) or treatment of other conditions (e.g. microalbuminuria). Conversely, the use of a single ICD-10 diagnosis code may overestimate the cohort with the diagnosis of interest; future longitudinal analyses using two ICD diagnoses (2 outpatient or 1 inpatient and 1 outpatient) may improve our ability to discern amongst diabetes types. Additional studies of the DCR may assess the sensitivity and specificity of variable approaches to identifying diagnoses of interest. Data extracted from EHRs are subject to inaccuracy due to errors or missing data, including bias introduced by individuals who are not actively seeking care and are subject to poorer health outcomes.<sup>7, 16, 25, 26</sup> To reduce errors, the DCR uses proprietary data collection software to ensure systematic data collection from participating practices, and data are subject to audits by CMS-approved third-party entities. NHANES is also subject to selection bias from lower participation rates, but the impact of this remains uncertain.<sup>27</sup> Data completeness for key variables in the DCR and NHANES cohorts is presented in Supplemental Table 1.

Treatment guidelines for both adults with type 1 and type 2 diabetes emphasize the importance of attaining metabolic control to reduce the risk of complication. It is likely that some differences (e.g. insulin use) are present between adults with different forms of diabetes. We chose to include adults with either type 1 or type 2 diabetes given the inherent challenges in accurately discriminating between the two diagnoses using either EHR (for the DCR) or NHANES data. While ICD codes for diabetes are generally accurate, they may not be reliable for type 1 or type 2 diabetes. Similarly, adjunctive use of medications, such as metformin or GLP-1 agonists, in the management of type 1 diabetes prevents reliable discrimination between these diagnoses based on medication use. The percentage of type 1 diabetes is comparably low in these large cohorts and the analyses best represent the care of adults with type 2 diabetes. There is an effort to define additional data for the DCR which may improve our ability to discern type 1 and type 2 diabetes, and we would recommend a similar approach to the NHANES dataset. Future analyses, using either longitudinal data or a broader dataset, may permit separate analyses of adults with type 1 or type 2 diabetes.

## 5. Conclusion

The DCR is the largest outpatient diabetes registry in the US collecting structured data from over 2900 ambulatory practices. Registry analyses may inform longitudinal trends in diabetes care and health outcomes at both the national and regional level. Further expansion of the registry cohort, especially in the West, and application of advanced sample weights may permit serial, nationally-representative, regional analyses of diabetes care delivery and related outcomes in the primary and specialty care settings to supplement national evaluations, such as NHANES. These analyses can support health policy initiatives and

efforts to cost-effectively manage the diabetes epidemic confronting the US.  
2, 4, 5, 10, 11, 28–31

## Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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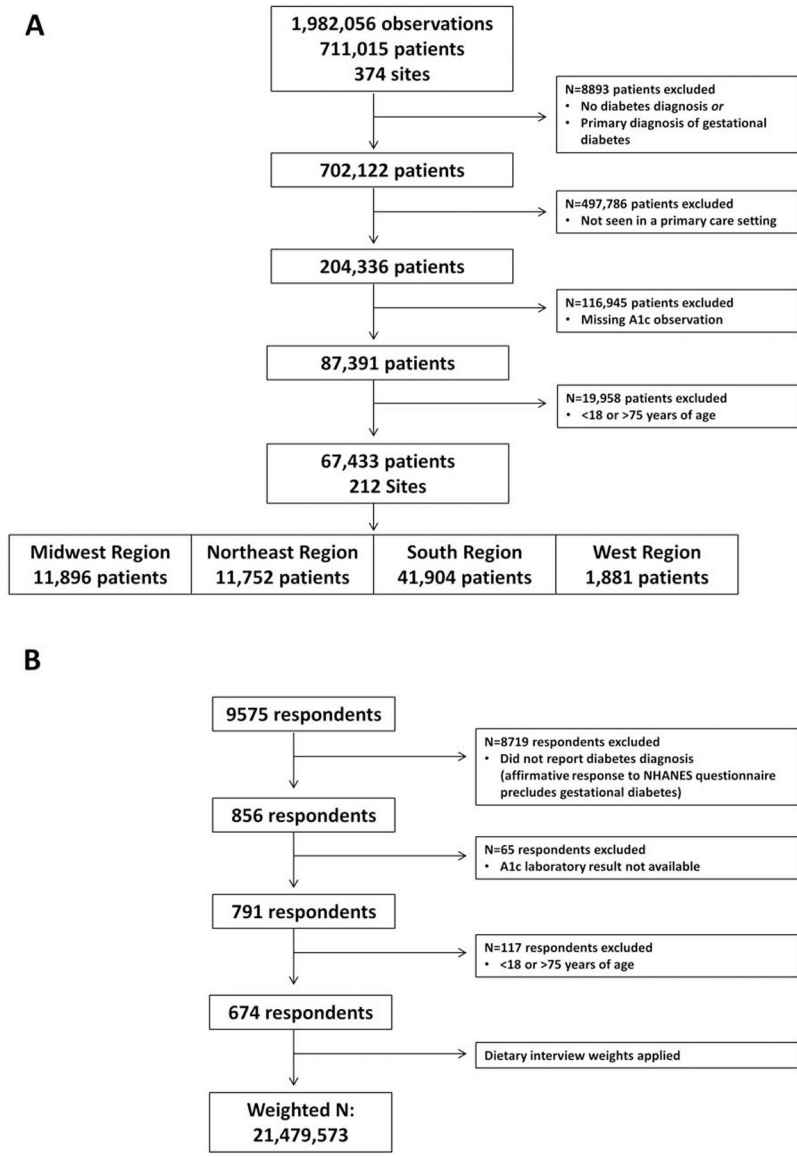
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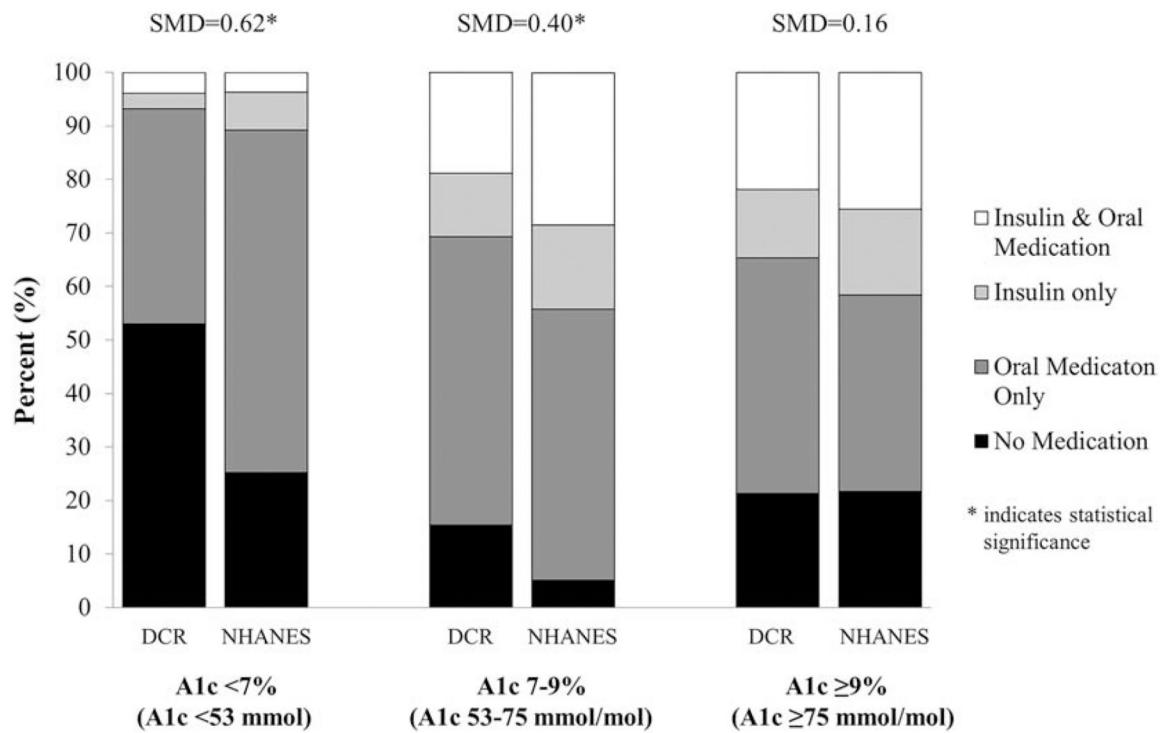
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### Highlights

- In the US, the diagnosis and effective management of diabetes and cardiovascular disease (CVD) risk factors in persons with diabetes remains suboptimal at the national level.
- There is significant regional variation in the diagnosis and management of diabetes and CVD risk factors which relate to regional differences in CVD risk factor control.
- Regional evaluations of diabetes care delivery are needed to inform more local, cost-effective disease management strategies for persons with diabetes.



**Figure 1.** Sample derivation for the (A) DCR and (B) NHANES cohorts.



**Figure 2.**

Diabetes management—no medications (N), oral agents only (O), insulin only (I) or insulin and oral agents (I+) —in DCR and NHANES by A1c level. For A1c<7%/<53 mmol/mol: DCR (N=53%, O=40%, I=3%, I+ =4%) vs. NHANES (N=25%, O=64%, I=7%, I+ =4%). For A1c 7–9%/53–75 mmol/mol: DCR (N=15%, O=54%, I=12%, I+ =19%) vs. NHANES (N=5%, O=51%, I=16%, I+ =28%). For A1c>9%/>75 mmol/mol: DCR (N=21%, O=44%, I=13%, I+ =22%) vs. NHANES (N=22%, O=37%, I=16%, I+ =26%).

**Table 1.**

Demographic and clinical characteristics of the DCR and NHANES cohorts.

	DCR N=67433	NHANES N=21479573*	p value	SMD <sup>†</sup>
Age (years)	61 ± 10	57 ± 12	<0.001	0.38
Sex (male)	52	56	<0.001	0.08
Race				
White non-Hispanic	83	76	<0.001	0.30
Black non-Hispanic	14	15		
Other	3	9		
Insurance coverage	100	91	<0.001	0.42
Height (cm)	170 ± 11	168 ± 10	0.08	0.18
Weight (kg)	97 ± 24	94 ± 23	0.22	0.12
Body mass index (kg/m <sup>2</sup> )	33.3 ± 7.3	33.2 ± 7.6	0.96	0.05
Body mass index category				
Underweight (<18.5 kg/m <sup>2</sup> )	0.5	0.5	<0.001	0.03
Normal (18.5–24.9 kg/m <sup>2</sup> )	9	10		
Overweight (25–29.9 kg/m <sup>2</sup> )	26	26		
Obese (≥ 30 kg/m <sup>2</sup> )	65	64		
HbA1c (%)	7.5 ± 2.2	7.4 ± 1.8	0.60	0.03
HbA1c (mmol/mol)	58.1 ± 24.4	57.3 ± 20.0	0.60	0.03
HbA1c category			<0.001	0.16
<7% (<53 mmol/mol)	56	51		
7–9% (53–75 mmol/mol)	27	35		
>9% (>75 mmol/mol)	16	15		
Systolic blood pressure (mmHg)	130 ± 17	129 ± 19	0.29	0.08
Diastolic blood pressure (mmHg)	76 ± 10	70 ± 13	<0.001	0.58
Blood pressure < 140/90 mmHG	70	74	<0.001	0.09
Total cholesterol (mmol/L)	4.45 ± 1.09	4.66 ± 1.17	0.050	0.18
High density lipoprotein (mmol/L)	1.22 ± 0.39	1.24 ± 0.39	0.73	0.03
Low density lipoprotein (mmol/L)	2.43 ± 0.91	3.34 ± 2.38	<0.001	0.50
Triglycerides (mmol/L)	1.81 ± 1.07	2.21 ± 1.63	0.007	0.29
Low density lipoprotein < 2.59 mmol/L	61	41	<0.001	0.39
'ABC' control <sup>††</sup>	18.8	6.0	<0.001	0.40
Tobacco use	17	32	<0.001	0.35
Hypertension	81	66	<0.001	0.37
Dyslipidaemia	83	63	<0.001	0.48
Medication Use				
Insulin	18	27	<0.001	0.22
Oral hypoglycaemic agent	56	71	<0.001	0.33

	<b>DCR</b> <b>N=67433</b>	<b>NHANES</b> <b>N=21479573*</b>	<b>p value</b>	<b>SMD<sup>†</sup></b>
	<b>Mean ± SD / %</b>	<b>Mean ± SD / %</b>		
Cholesterol-lowering	71	67	<0.001	0.08
Anti-hypertensive	79	95	<0.001	0.50

\* Weighted sample size

<sup>†</sup> Standardized mean difference

<sup>††</sup> HbA1c < 7% (< 53 mmol/mol), blood pressure < 140/90 mmHg and low density lipoprotein < 2.59 mmol/L

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**Table 2.**

Demographic and clinical characteristics of the DCR cohort by geographical region.

	Midwest N=11896	Northeast N=11753	South N=41904	West N=1881	p value
	Mean ± SD / %	Mean ± SD / %	Mean ± SD / %	Mean ± SD / %	
Age (years)	62 ± 10	61 ± 10	61 ± 11	64 ± 9	<0.001
Sex (male)	53	56	50	59	<0.001
Race					<0.001
White non-Hispanic	85	89	81	88	
Black non-Hispanic	13	8	17	6	
Other	2	3	2	7	
Insurance coverage	99	100	100	99	<0.001
Height (cm)	170 ± 11	170 ± 11	170 ± 11	171 ± 11	<0.001
Weight (kg)	100 ± 25	94 ± 22	97 ± 23	98 ± 24	<0.001
Body mass index (kg/m <sup>2</sup> )	34.4 ± 7.9	32.3 ± 6.8	33.3 ± 7.3	33.2 ± 7.2	<0.001
Body mass index category					<0.001
Underweight (<18.5 kg/m <sup>2</sup> )	0.5	0.5	0.5	0.5	
Normal (18.5–24.9 kg/m <sup>2</sup> )	8	11	9	9	
Overweight (25–29.9 kg/m <sup>2</sup> )	22	29	26	24	
Obese (≥ 30 kg/m <sup>2</sup> )	70	60	65	66	
HbA1c (%)	8.5 ± 3.0	7.0 ± 1.7	7.3 ± 2.1	7.5 ± 1.9	<0.001
HbA1c (mmol/mol)	68.9 ± 32.5	52.5 ± 18.1	56.5 ± 22.5	58.2 ± 20.9	<0.001
HbA1c category					<0.001
<7% (<53 mmol/mol)	43	66	58	51	
7–9% (53–75 mmol/mol)	28	25	27	34	
>9% (>75 mmol/mol)	29	10	15	16	
Systolic blood pressure (mmHg)	129 ± 18	128 ± 15	131 ± 17	128 ± 17	<0.001
Diastolic blood pressure (mmHg)	73 ± 12	77 ± 9	77 ± 10	74 ± 11	<0.001
Blood pressure < 140/90 mmHG	72	74	68	73	<0.001
Total cholesterol (mmol/L)	4.22 ± 1.09	4.55 ± 1.03	4.50 ± 1.09	4.11 ± 1.11	<0.001
High density lipoprotein (mmol/L)	1.22 ± 0.39	1.29 ± 0.41	1.22 ± 0.39	1.16 ± 0.36	<0.001
Low density lipoprotein (mmol/L)	2.20 ± 0.91	2.51 ± 0.88	2.51 ± 0.91	2.12 ± 0.88	<0.001
Triglycerides (mmol/L)	1.84 ± 1.04	1.75 ± 1.02	1.81 ± 1.10	1.85 ± 1.10	<0.001
Low density lipoprotein < 2.59 mmol/L	72	58	58	74	<0.001
'ABC' control †	14.3	20.9	19.5	18.7	<0.001
Tobacco use	27	13	16	19	<0.001
Hypertension	77	79	83	88	<0.001
Dyslipidaemia	76	85	85	89	<0.001
Medication Use					
Insulin	27	15	16	33	<0.001
Oral hypoglycaemic agent	59	50	55	69	<0.001
Cholesterol-lowering	77	67	69	88	<0.001

	<b>Midwest</b> N=11896	<b>Northeast</b> N=11753	<b>South</b> N=41904	<b>West</b> N=1881	
	<b>Mean ± SD / %</b>	<b>Mean ± SD / %</b>	<b>Mean ± SD / %</b>	<b>Mean ± SD / %</b>	<b>p value</b>
Anti-hypertensive	89	72	78	96	<0.001

<sup>†</sup>HbA1c<7% (<53 mmol/mol), blood pressure <140/90 mmHg and low density lipoprotein <2.59 mmol/L

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