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## Intensive Glycemic Control in Younger and Older U.S. Adults with Type 2 Diabetes

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### Abstract

**Aims**—To determine the extent to which older vs. younger adults with diabetes intensively control glycemia.

**Methods**—Participants were age 40 years who self-reported a physician diagnosis of diabetes in the 2009–2014 National Health and Nutrition Examination Surveys (N=1,554). Intensive glycemic control was defined as A1c<7.0% and taking insulin, sulfonylureas, or 2 glycemic medications. Logistic regression was used to determine the adjusted odds of intensive control in older (65 years) vs. younger adults (age 40–64 years).

**Results**—The prevalence of intensive control was greater for older (33.4%) vs. younger (21.3%) adults (p<0.001). In logistic regression, intensive control was significantly higher in older vs. younger adults after fully adjusting for sociodemographics, diabetes duration, comorbidities, disability, use of multiple medications, and depression (OR=1.72, 1.09–2.69). The multivariable adjusted prevalence of intensive control was 40% higher in adults 75 years (35.6%) compared to adults 40–49 years (21.7%).

**Conclusions**—Older adults are being treated more aggressively than younger adults to achieve A1c<7.0% despite the presence of comorbidities, duration of diabetes, disability, and depression. Glycemic guidelines for individualized therapy are not being widely followed.

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**Conflicts of Interest:** None

## Keywords

Diabetes; Treatment by age; NHANES

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## 1. Introduction

An estimated 28.9 million U.S. adults have diabetes, yielding a large public health burden of morbidity, mortality and economic costs due mainly to diabetes-related complications<sup>1</sup>. The Diabetes Control and Complications Trial (DCCT) and the United Kingdom Prospective Diabetes Study (UKPDS) established that intensive glycemic control with a reduction in A1c levels to an average of 7.0% significantly reduced microvascular disease in persons with type 1 and type 2 diabetes<sup>2,3</sup>. Clinical practice guidelines developed on the basis of these findings recommended an A1c<7.0% to decrease the risk of diabetes complications. After the trials ended, significant long term reductions in cardiovascular disease (CVD) in those randomized to intensive treatment emerged during participant follow-up<sup>4</sup>. However, three subsequent studies, in older adults with longer duration type 2 diabetes and CVD or risk factors for CVD, found that more intensive therapy targeting even lower A1c levels did not reduce CVD<sup>5-7</sup>. Moreover, the Action to Control Cardiovascular Risk in Diabetes (ACCORD) trial, which targeted A1c<6.0%, increased mortality compared to the standard (A1c<7.5%) treatment group<sup>6</sup>.

On the basis of these studies, the ADA revised their guidelines in 2008 to recommend an A1c goal of <7.0% for adults who can benefit the most from a reduction in A1c to prevent diabetes-related complications, such as those with longer life expectancy and little comorbidity, but less stringent goals (e.g., A1c <8.0%) for patients with a history of hypoglycemia, advanced complications, several comorbid conditions, and shorter life expectancy<sup>8</sup>. Similarly, the American Geriatrics Society (AGS) first recommended in 2003 that individualized therapy take into account diabetes severity and life expectancy and recommended less stringent targets (e.g., A1c <8.0%) when the risks of intensive glycemic control outweighed the benefits<sup>9</sup>.

A previous study using national data from 2001–2010 found that among older adults with A1c<7.0% and significant health problems, 60% were treated with insulin or sulfonylureas; these results indicate possible overtreatment<sup>10</sup>. However, the practice patterns in younger adults with longer life expectancy, versus older adults, are relatively unknown. We used data from the National Health and Nutrition Examination Survey to determine the extent to which younger versus older adults are being treated more intensively to lower A1c levels while accounting for factors related to treatment, including duration of disease, comorbidities, disability, use of prescription medications, and depression.

## 2. Subjects, Materials, and Methods

### 2.1 Study Design and Participants

The National Health and Nutrition Examination Survey (NHANES) is a stratified multistage probability cluster survey conducted in the non-institutionalized civilian U.S. population<sup>11</sup>.

Participants are interviewed in their home for demographic and health information and are then scheduled to visit a mobile examination center for physical examinations and laboratory measures<sup>12,13</sup>. Written informed consent was obtained from all participants and was approved by the National Center for Health Statistics Institutional Review Board. Our analyses included adults age  $\geq 40$  years who answered “yes” when asked whether a physician or other health care professional ever told them they had diabetes (N=1,554). We excluded adults with probable type 1 diabetes defined as having a diagnosis at age  $<30$  years, starting insulin treatment within one year of diagnosis, and currently taking insulin (n=21). Participants self-reported age, race/ethnicity, education, health insurance status, smoking status, duration of diabetes, and use of insulin.

## 2.2 Health Status

Self-reported comorbidities included a history of cancer (excluding skin cancer, except Melanoma), lung disease (asthma, bronchitis, or emphysema), cardiovascular disease (stroke, congestive heart failure, coronary heart disease, angina, or heart attack). Chronic kidney disease (CKD) was determined using the Chronic Kidney Disease Epidemiology Collaboration (CKD-EPI) equation which estimates glomerular filtration rate (eGFR) from serum creatinine based on age, sex, and race and defined as  $eGFR < 60 \text{ mL/min per } 1.73 \text{ m}^2$ <sup>14</sup>. Participants with  $\geq 1$  of these conditions were considered to have comorbidities.

Disability was defined as having mobility disability, work disability, or general pain. Participants who reported needing special equipment to walk or much difficulty/inability to do any of the following activities were considered to have a mobility disability: (1) walking a quarter mile, (2) walking up ten steps, (3) stooping, crouching, or kneeling, (4) walking between rooms, (5) standing up from an armless chair, (6) getting in and out of bed. A work disability was determined if participants responded “yes” when asked if limitations kept them from working. Participants who reported  $\geq 3$  days in the past month where pain made it hard for usual activities were considered to have disability related to general pain.

Participants self-reported use of prescription medications in the past 30 days and were asked to show the interviewer their medication containers. Use of  $\geq 6$  prescription medications was considered multiple medication use.

The Patient Health Questionnaire (PHQ9) was used to determine depression<sup>15</sup>. Participants answered 9 questionnaire items with “not at all” (value of 0), “several days” (value of 1), “more than half the days” (value of 2), or “nearly every day” (value of 3) in the past two weeks.” Symptoms included (1) little interest in doing things, (2) feeling down, depressed, or hopeless, (3) trouble sleeping or sleeping too much, (4) feeling tired or having little energy, (5) poor appetite or overeating, (6) feeling bad about yourself, (7) trouble concentrating on things, (8) moving or speaking slowly or too fast, (9) would be better off dead. Depression was defined as having a PHQ9 score  $\geq 10$ .

## 2.3 Intensive Control of Diabetes

Intensive control of diabetes was defined as  $A1c < 7.0\%$  and use of sulfonylureas, insulin, or  $\geq 2$  glycemic medications. The DCCT used insulin and the UKPDS used insulin and sulfonylureas as intensive therapy to achieve  $A1c$  control<sup>2,3</sup> and taking 2 or more glycemic

medications indicates that diabetes cannot be controlled with lifestyle or first line medications. This definition of intensive control is not based on A1c alone but in the context of pharmacological therapy (and potential detriment) required to achieve near normal A1c. The comparison in older and younger adults is the use of this potentially harmful pharmacological therapy to achieve A1c<7.0%. Hemoglobin A1c was measured in all adults from a standard blood draw and standardized to the Diabetes Control and Complications Trial method<sup>16</sup>. A1c was measured with the A1c G7 HPLC Glycohemoglobin Analyzer (Tosoh Medics, Inc., San Francisco, CA) which had a coefficient of variation of 0.7–1.5%. Sulfonylurea use was determined during the prescription medication interview; insulin use was self-reported and adjudicated with the prescription medication interview.

## 2.4 Statistical Analysis

Distributions of demographic factors, health insurance, smoking status, and duration of diabetes and prevalences of comorbidities, disability, use of multiple medications, and depression were determined by age (40–64 years vs. ≥65 years). Analysis of variance (F-test) was used to determine differences in the distribution of participant characteristics. Prevalence of intensive control of diabetes was determined by age and by demographics, presence of comorbidities, disability, use of multiple medications, and depression. Differences in means were tested by two-tailed large sample z-tests. Logistic regression (odds ratio [OR], 95% confidence interval [CI]) was used to determine the odds of older vs. younger adults having intensive control of diabetes. Models were (1) unadjusted, (2) adjusted for sex and race/ethnicity, (3) additionally adjusted for health insurance and education, (4) additionally adjusted for duration of diabetes, (5) additionally adjusted for smoking, (6) additionally adjusted for comorbidities, disability, and use of multiple medications, (9) and additionally adjusted for depression. Predictive margins regression was used to determine the prevalence of intensive control by smaller age groups (40–49, 50–64, 65–74, ≥75 years); models were adjusted for covariates in the same order as for the logistic regression analysis. These regression techniques account for differences in participant characteristics (e.g. comorbidities, duration of diabetes), by averaging these characteristics across all subgroups and creating age groups with participants who have similar levels of comorbidities, duration of diabetes, disability, medication use, and depression. All statistical analyses used sample weights and accounted for the cluster design using SUDAAN (SUDAAN User's Manual, Release 9.2, 2008; Research Triangle Institute).

## 3. Results

### 3.1 Characteristics of the Study Population

Fifty-five percent (55.2%) were age 40–64 years (median age 54.4 years) and 44.8% were ≥65 years (median age 72.1 years); 48.8% were women with no difference by age group (Table 1). The majority was non-Hispanic white (61.4%), and most had a high school or more education (73.4%). Almost all adults with type 2 diabetes age ≥65 years had health insurance (98.1%) compared to 85.1% of those age 40–64 years ( $p<0.001$ ). More adults age 40–64 years were current smokers compared to those age ≥65 years ( $p<0.001$ ). Duration of diabetes ≥20 years was higher for adults age ≥65 years compared to 40–64 years ( $p<0.001$ ). The prevalence of comorbidities, disability, and use of multiple medications was greater for

those age  $\geq 65$  years vs. 40–64 years ( $p = 0.006$  for all). More adults age 40–64 years vs.  $\geq 65$  years reported depression ( $p < 0.001$ ).

### 3.2 Prevalence of Intensive Control of Diabetes

The prevalence of intensive control was higher for adults age  $\geq 65$  years vs. age 40–64 years ( $p < 0.001$ ) (Table 2). This relationship by age was consistent for women, all race/ethnicities, and education levels; for those with health insurance; for never or former smokers; and for those with duration of diabetes of 5–19 years ( $p < 0.05$  for all).

The prevalence of intensive control of diabetes was higher among those age  $\geq 65$  years with comorbidities compared to those age 40–64 years with comorbidities ( $p = 0.018$ ). In addition, prevalence of intensive control was higher among adults age  $\geq 65$  years with disability ( $p < 0.001$ ), or using multiple medications ( $p = 0.001$ ), compared to their counterparts age 40–64 years.

### 3.3 Logistic Regression Analysis

Adults age  $\geq 65$  years were significantly more likely to use intensive control compared to those age 40–64 years after adjusting for demographics, duration of diabetes, smoking, comorbidities, disability, use of multiple medications, and depression (OR=1.72, 1.09–2.69) (Table 3). Thus, regardless of the differences in health status by age, these results indicate that older adults were more likely to have intensive glycemic control compared to younger adults. The adjusted association was slightly attenuated from the unadjusted model.

### 3.4 Adjusted Prevalence of Intensive Control of Diabetes

The prevalence of intensive control increased with increasing age; unadjusted estimates and estimates adjusted for sex and race/ethnicity were similar (Figure 1, Appendix Table A). The magnitude of intensive control was slightly attenuated for older adults in the fully adjusted model but still remained significantly higher compared to younger adults. In the fully adjusted model, the prevalence of intensive control was 21.7%, 23.5%, 32.5%, and 35.6% for those age 40–49, 50–64, 65–74, and  $\geq 75$  years, respectively.

## 4. Discussion

Nearly 26 percent of people age  $\geq 65$  years had diabetes in 2012, with a substantial increase in the number of older adults with diabetes expected in the coming decades<sup>1</sup>. There is considerable heterogeneity in this group. The older population with diabetes includes individuals newly diagnosed with diabetes and as well as those with longer durations of disease who may have established complications. While diabetes in older adults is associated with increased mortality and reduced functional status, the population also includes those with substantial life expectancy and without comorbid conditions. Thus, recommendations for glycemic targets are not age specific but rather depend on individualized assessments of the risks and benefits of treatment: balancing factors such as life expectancy of sufficient duration to realize reduction in microvascular complications with glycemic control and risk of hypoglycemia and other harms of glucose lowering medications.

While age may not be the major determinant of life expectancy for individuals, in the population as a whole, age does strongly correlate with life expectancy. Thus, it would be expected that population-wide the implementation of individualized goals for therapy, in which life expectancy is a major factor, would lead to less intensive control on average in older versus younger populations with diabetes. We found that, regardless of demographics, duration of diabetes, comorbidities, and other diabetes-related factors, older adults were more likely to have intensive glycemic control compared to their younger counterparts. Moreover, there was roughly a 40% increase in the adjusted prevalence of intensive glycemic control for adults age ≥ 75 years vs. age 40–49 years. Thus, despite treatment recommendations by the ADA and AGS which endorse less stringent goals for patients with shorter life expectancy and comorbidities,<sup>17,18</sup> older adults are being treated more aggressively than younger adults.

Our results agree with a recent national study which found that adults age ≥ 65 years were achieving tight glycemic targets regardless of poor health status and that most of these patients were treated with insulin or sulfonylureas<sup>10</sup>. However, that study did not compare younger and older populations with diabetes and was unable to assess any response to the change in ADA Guidelines after the ACCORD results were released.

Recent data indicate that hypoglycemia has surpassed hyperglycemia as a cause of emergency room visits and hospitalizations<sup>19</sup>. About 25% of hospitalizations for adverse drug events in U.S. adults age ≥ 65 years are due to insulin or oral hypoglycemic agents<sup>20</sup>. Thus, diabetes agents were selected as one of the top three concerns in the recent National Action Plan for Adverse Drug Events<sup>21</sup>. The ADA, AGS, and the Department of Veterans Affairs/Department of Defense all recommend that risk of hypoglycemia be considered in recommending a target A1c goal<sup>8,9,22</sup>. Older adults may be particularly vulnerable to hypoglycemia<sup>23</sup>. After adjustment for duration of diabetes, rates of hypoglycemia increase sharply with advancing age<sup>24</sup>. Some individuals, particularly early in the course of diabetes, may achieve tight glucose control with low risk therapies such as lifestyle modifications or metformin. However, our definition of intensive control required use of sulfonylurea, insulin or at least two anti-diabetic agents. We found that despite their increased risk of hypoglycemia older adults were more likely to receive intensive glycemic control compared to younger adults, even after adjustment for duration of diabetes.

Several factors may contribute to more intensive treatment of older adults even after controlling for diabetes-related factors. First, older individuals may have more interaction with doctors due to multiple medical problems. Second, older adults generally have more time to take care of their health and/or are more concerned about their health than younger adults. Third, although our finding of more intensive control was not altered by adjustment for insurance status, Medicare coverage may lessen barriers to intensive control compared to insurance with more out of pocket costs. Finally, diabetes in older adults may be less complicated and easier to treat due to survival bias<sup>25</sup>.

One strength of our study is the use of nationally representative data allowing generalization to the U.S. adult non-institutionalized civilian population. The NHANES provides standardized measures of A1c to characterize, in part, intensive glycemic control. The

NHANES included several covariates, including duration of diabetes, comorbidities, disability, prescription medication use, and depression, which allowed us to adjust our analysis for these confounding factors. One limitation is the reliance on self-reported medication use and the lack of information on medication adherence. In addition, information on the length of time participants were on intensive treatment to achieve  $A1c < 7.0\%$  was not available.

#### 4.1 Conclusions

Guidelines for glycemic control call for individualized targets that weigh the risks and benefits of intensive therapy. While age is not a suitable basis for therapeutic decision-making in individual patients, in the population at large it would be expected to correlate with health status. Indeed, we found that older adults had more comorbidities, disability, and use of multiple medications than younger adults and, even after controlling for these factors, older adults were treated more aggressively than younger adults with agents that carry the risk of hypoglycemia. This suggests that the guidelines for individualized therapy are not being widely followed.

#### Acknowledgments

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None

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## Appendix

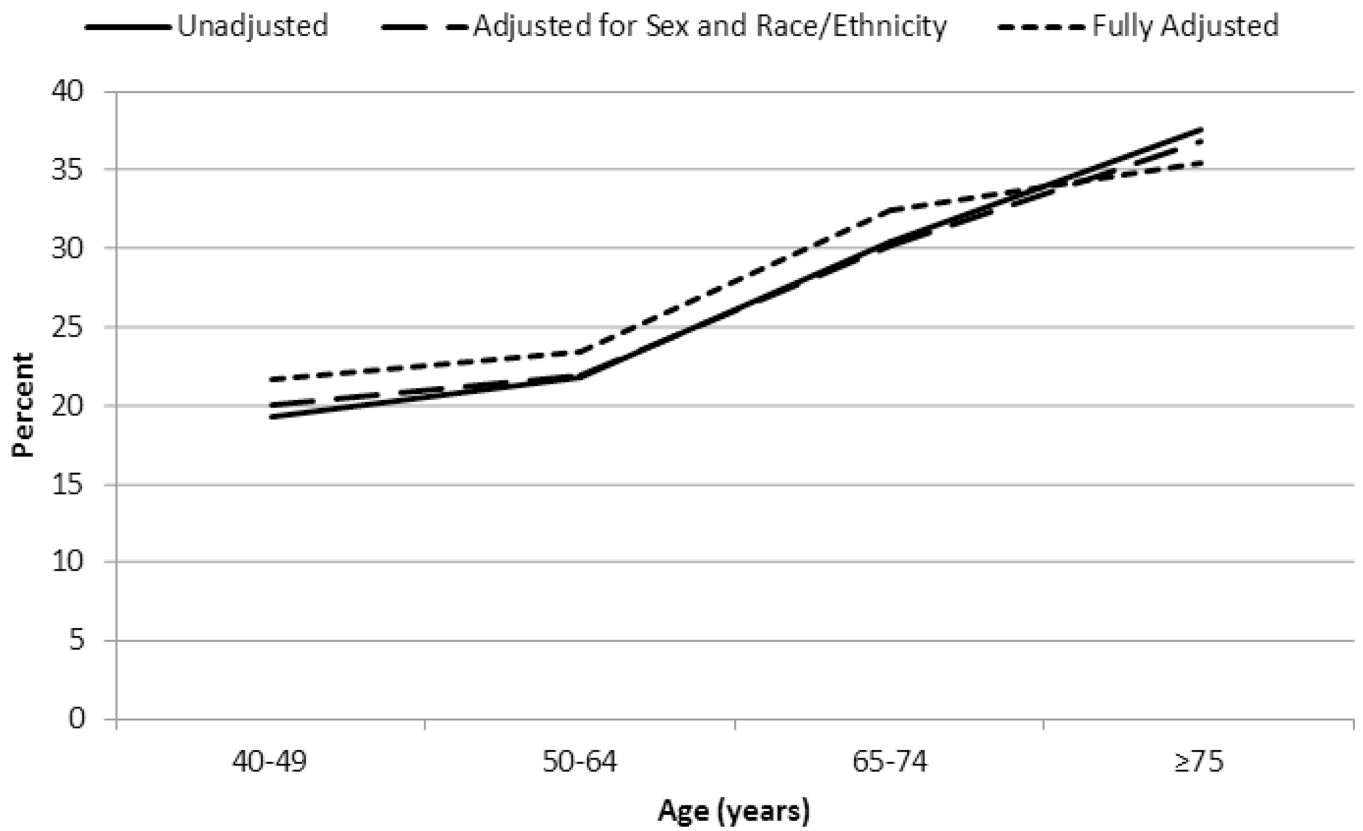
**Table A**

Prevalence (standard error) of intensive glycemic control adjusted for covariates among adults with type 2 diabetes, NHANES 2009–2014

	Model 1	Model 2	Model 3	Model 4	Model 5	Model 6	Model 7
	Unadjusted	Adjusted for Sex, Race/Ethnicity	Model 2+ Health Insurance, Education	Model 3+ Duration of Diabetes	Model 4+ Smoking	Model 5+ Comorbidities, Disability, Multiple Medication Use*	Model 6 + Depression <sup>†</sup>
<b>Age</b>							
40–49	19.3 (3.46)	20.1 (3.71)	21.3 (3.84)	21.3 (3.85)	21.4 (3.91)	20.1 (3.60)	21.7 (3.89)
50–64	21.9 (2.50)	22.0 (2.52)	22.6 (2.58)	22.5 (2.55)	22.8 (2.61)	23.8 (3.04)	23.5 (3.42)
65–74	30.5 (2.88)	30.3 (2.83)	29.6 (2.67)	29.8 (2.63)	29.5 (2.63)	33.0 (3.12)	32.5 (3.24)
75	37.7 (3.31)	36.9 (3.30)	35.0 (3.27)	34.5 (3.37)	34.1 (3.53)	36.1 (3.99)	35.6 (4.29)

\* Comorbidities include cancer (excluding skin cancer except Melanoma), lung disease (asthma, bronchitis, or emphysema), cardiovascular disease (stroke or CVD), chronic kidney disease (CKD-EPI equation); Disability includes work disability, mobility disability, or pain 3 days in the past month that makes daily activities difficult; Use of 6 prescription medications

<sup>†</sup> Depression defined as a PHQ9 score  $\geq 10$



**Figure 1.** Prevalence of intensive glycemic control by age among adults with type 2 diabetes, NHANES 2009–2014. Fully adjusted model includes sex, race/ethnicity, health insurance, education, duration of diabetes, smoking, comorbidities, disability, multiple medication use, and depression.

**Table 1**

Characteristics of adults with type 2 diabetes age ≥ 40 years, NHANES 2009–2014

	Percent (standard error)			ANOVA p-value
	Total (N=1,554)	Age 40–64 Years (n=793)	Age ≥ 65 Years (n=761)	
<b>Age</b>	100.0	55.2 (1.51)	44.8 (1.51)	<b>0.005</b>
<b>Sex, % women</b>	48.8 (1.67)	49.2 (2.28)	48.2 (2.61)	<b>0.778</b>
<b>Race/Ethnicity</b>				<b>&lt;0.001</b>
Non-Hispanic white	61.4 (2.31)	57.1 (2.87)	66.8 (2.49)	
Non-Hispanic black	15.8 (1.67)	17.8 (2.07)	13.3 (1.49)	
Mexican American	8.6 (1.55)	11.0 (1.88)	5.6 (1.37)	
Other Hispanic	5.0 (0.77)	5.5 (0.93)	4.4 (0.82)	
Non-Hispanic Other	9.2 (1.27)	8.6 (1.30)	10.0 (1.67)	
<b>Education, % high school graduate or higher</b>	73.4 (1.89)	75.7 (2.14)	70.6 (2.30)	<b>0.029</b>
<b>Health Insurance, % yes</b>	90.9 (0.97)	85.1 (1.67)	98.1 (0.63)	<b>&lt;0.001</b>
<b>Current Smoking</b>	14.0 (0.92)	20.6 (1.52)	5.8 (0.84)	<b>&lt;0.001</b>
<b>Duration of Diabetes (years)</b>				<b>&lt;0.001</b>
<5	27.9 (1.42)	34.9 (2.05)	19.3 (1.88)	
5–9	23.0 (1.19)	24.6 (1.91)	20.9 (1.44)	
10–19	31.2 (1.91)	30.0 (2.48)	32.7 (2.06)	
≥ 20	17.9 (1.18)	10.5 (1.40)	27.1 (1.88)	
<b>Comorbidities, 1<sup>*</sup></b>	60.4 (1.71)	47.5 (2.20)	76.7 (1.73)	<b>&lt;0.001</b>
<b>Disability<sup>†</sup></b>	67.4 (2.25)	63.2 (2.95)	72.3 (2.48)	<b>0.006</b>
<b>Multiple Medication Use, % ≥ 6 medications</b>	56.6 (1.97)	49.0 (2.65)	66.0 (2.04)	<b>&lt;0.001</b>
<b>Depression<sup>‡</sup></b>	12.4 (1.18)	16.2 (1.72)	7.8 (1.34)	<b>&lt;0.001</b>

\* Comorbidities include cancer (excluding skin cancer except Melanoma), lung disease (asthma, bronchitis, or emphysema), cardiovascular disease (stroke or CVD), and chronic kidney disease

<sup>†</sup> Disability includes work disability, mobility disability, or pain ≥ 3 days in past month that makes daily activities difficult

<sup>‡</sup> Depression defined as a PHQ9 score ≥ 10

**Table 2**

Prevalence of intensive glycemic control of type 2 diabetes among adults age ≥40 years, NHANES 2009–2014

	Percent (standard error)			p-value
	Total (N=1,554)	Age 40–64 Years (n=793)	Age ≥65 Years (n=761)	
<b>Age</b>				
Total	26.7 (1.59)	21.3 (2.32)	33.4 (2.18)	<0.001
<b>Sex</b>				
Men	27.7 (2.07)	23.2 (3.02)	33.1 (3.53)	0.057
Women	25.7 (2.03)	19.3 (2.64)	33.7 (2.39)	<0.001
<b>Race/Ethnicity</b>				
Non-Hispanic white	29.1 (2.29)	22.7 (3.52)	35.8 (3.01)	0.007
Non-Hispanic black	27.7 (2.52)	23.9 (2.82)	34.1 (4.28)	0.042
Mexican American	16.5 (1.90)	13.2 (2.49)	24.7 (4.00)	0.032
Other Hispanic	17.3 (3.23)	12.4 (3.33)	24.7 (4.92)	0.034
<b>Education</b>				
<HS	31.6 (2.41)	23.8 (3.46)	39.5 (3.21)	0.001
HS	24.9 (1.97)	20.5 (2.71)	30.9 (2.84)	0.011
<b>Health Insurance</b>				
No	13.5 (3.93)	13.6 (4.20)	12.7 (7.08)	0.914
Yes	28.1 (1.73)	22.6 (2.49)	33.9 (2.30)	0.001
<b>Smoking</b>				
Current	21.8 (3.62)	20.0 (4.06)	29.7 (7.81)	0.267
Never/former	27.5 (1.78)	21.6 (2.71)	33.7 (2.38)	0.002
<b>Duration of Diabetes (years)</b>				
<5	26.0 (2.88)	26.1 (4.01)	25.8 (3.55)	0.963
5–9	26.6 (3.23)	18.9 (3.89)	37.9 (5.76)	0.010
10–19	23.6 (2.30)	16.5 (2.60)	31.8 (3.91)	0.002
≥20	32.9 (3.52)	23.1 (6.23)	37.6 (4.05)	0.054
<b>Comorbidities 1*</b>				
No	21.0 (2.23)	17.9 (2.90)	29.7 (4.85)	0.067
Yes	30.7 (2.21)	24.8 (3.41)	35.3 (2.75)	0.018
<b>Cancer</b>				
No	25.8 (1.79)	21.1 (2.57)	32.4 (2.29)	0.002
Yes	32.4 (4.44)	24.5 (8.93)	37.7 (4.62)	0.203
<b>Lung Disease</b>				
No	25.8 (2.00)	20.9 (2.86)	31.6 (2.37)	0.004
Yes	29.2 (2.92)	22.4 (4.03)	38.4 (4.93)	0.019
<b>Cardiovascular Disease</b>				
No	25.3 (1.71)	21.1 (2.54)	32.4 (2.97)	0.013

	Percent (standard error)			p-value
	Total (N=1,554)	Age 40–64 Years (n=793)	Age 65 Years (n=761)	
Yes	30.3 (3.45)	22.4 (4.48)	35.1 (3.80)	<b>0.008</b>
Chronic Kidney Disease				
No	24.0 (1.72)	20.0 (2.28)	31.7 (3.25)	<b>0.009</b>
Yes	35.8 (3.32)	31.5 (6.35)	37.1 (3.65)	0.426
<b>Disability<sup>†</sup></b>				
No	26.7 (3.24)	21.6 (4.81)	34.6 (5.21)	0.098
Yes	29.4 (1.80)	21.7 (2.48)	37.3 (2.52)	<b>&lt;0.001</b>
Work Disability				
No	25.3 (2.08)	20.8 (3.17)	30.4 (2.79)	<b>0.032</b>
Yes	30.7 (2.78)	22.5 (2.25)	43.1 (4.92)	<b>&lt;0.001</b>
Mobility Disability				
No	25.5 (2.33)	22.4 (3.28)	30.3 (3.74)	0.147
Yes	28.8 (2.25)	18.6 (2.78)	37.4 (3.14)	<b>&lt;0.001</b>
Pain, general				
No	29.5 (2.73)	22.0 (4.51)	37.7 (3.96)	<b>0.019</b>
Yes	32.2 (2.50)	22.8 (4.02)	43.7 (3.89)	<b>0.002</b>
<b>Multiple Medication Use, ≥ 6 medications</b>				
No	20.4 (2.19)	19.2 (2.99)	22.7 (3.18)	0.454
Yes	31.5 (2.27)	23.4 (3.32)	39.0 (3.18)	<b>0.001</b>
<b>Number of Medications</b>				
1–3	16.1 (3.73)	15.8 (4.71)	17.0 (4.87)	0.848
4–5	23.6 (2.51)	22.1 (3.81)	25.9 (3.90)	0.533
6–9	29.1 (2.66)	22.7 (3.79)	34.8 (3.65)	<b>0.026</b>
10	37.2 (4.46)	24.9 (6.50)	49.6 (4.83)	<b>0.002</b>
<b>Depression (PHQ9 Score)<sup>‡</sup></b>				
<10	26.9 (1.69)	20.9 (2.67)	33.5 (2.48)	<b>0.003</b>
10	21.3 (2.84)	18.2 (3.24)	29.3 (6.64)	0.171

Intensive control of diabetes is defined as A1c<7.0% and taking insulin or sulfonylurea or ≥ 2 glycemic medications

\* Comorbidities include cancer (excluding skin cancer except Melanoma), lung disease (asthma, bronchitis, or emphysema), cardiovascular disease (stroke or CVD), and chronic kidney disease

<sup>†</sup>Disability includes work disability, mobility disability, or pain ≥ 3 days in past month that makes daily activities difficult

<sup>‡</sup>Depression defined as a PHQ9 score ≥ 10

**Table 3**

Odds ratio (95% confidence interval) for intensive glycemic control of type 2 diabetes, NHANES 2009–2014

	Model 1	Model 2	Model 3	Model 4	Model 5	Model 6	Model 7
	Unadjusted	Adjusted for Sex, Race/Ethnicity	Model 2 + Health Insurance, Education	Model 3+ Duration of Diabetes	Model 4+ Smoking	Model 5+ Comorbidity, Disability, Multiple Medication Use*	Model 6 + Depression <sup>†</sup>
Age (years)							
40–64	1.00	1.00	1.00	1.00	1.00	1.00	1.00
65	<b>1.86 (1.31, 2.63)</b>	<b>1.81 (1.26, 2.58)</b>	<b>1.64 (1.15, 2.33)</b>	<b>1.61 (1.12, 2.27)</b>	<b>1.59 (1.10, 2.32)</b>	<b>1.77 (1.22, 2.56)</b>	<b>1.72 (1.09, 2.69)</b>

\* Comorbidities include cancer (excluding skin cancer except Melanoma), lung disease (asthma, bronchitis, or emphysema), cardiovascular disease (stroke or CVD), and chronic kidney disease; Disability includes work disability, mobility disability, or pain > 3 days in past month that makes daily activities difficult; Use of > 6 prescription medications

<sup>†</sup> Depression defined as a PHQ9 score > 10