

### **HHS Public Access**

Author manuscript *J Pediatr*. Author manuscript; available in PMC 2015 December 23.

Published in final edited form as:

J Pediatr. 2009 November ; 155(5): 668-72.e1-3. doi:10.1016/j.jpeds.2009.05.025.

# Glycemic Control in Youth with Diabetes: The SEARCH for Diabetes in Youth Study

Diana B. Petitti, MD, MPH, Georgeanna J. Klingensmith, MD, Ronny A. Bell, PhD, MS, Jeanette S. Andrews, MS, Dana Dabelea, MD, PhD, Giuseppina Imperatore, MD, PhD, Santica Marcovina, PhD, Catherine Pihoker, MD, Debra Standiford, MSN, CNP, Beth Waitzfelder, PhD, and Elizabeth Mayer-Davis, PhD for the SEARCH for Diabetes in Youth Study Group<sup>\*</sup>

Biomedical Informatics, Arizona State University, Phoenix AZ (D.P.); Pediatric Clinics, Barbara Davis Center for Childhood Diabetes, University of Colorado at Denver and Health Sciences Center, Denver, CO (G.K.); Department of Epidemiology and Prevention, Wake Forest University School of Medicine, Winston-Salem, NC (R.B.); Department of Biostatistical Sciences, Division of Public Health Sciences, Wake Forest University Health Sciences, Winston-Salem, NC (J.A.); Department of Epidemiology, Colorado School of Public Health, University of Colorado, Denver, CO (D.D.); Division of Diabetes Translation, National Center for Chronic Disease Prevention and Health Promotion, Centers for Disease Control and Prevention, Atlanta, GA (G.I.); Department of Medicine, University of Washington, Seattle WA (S.M.); Division of Endocrinology and Diabetes, Children's Hospital and Regional Medical Center, Seattle WA (C.P.); Division of Endocrinology, Children's Hospital Medical Center, Cincinnati OH (D.S.); Pacific Health Research Institute, Honolulu HI (B.W.); Department of Nutrition, Gillings School of Global Public Health, University of North Carolina, Chapel Hill, NC (E.M.-D.); and Department of Epidemiology and Biostatistics, University of South Carolina, Columbia, SC (E.M.-D.)

#### Abstract

**Objective**—To assess correlates of glycemic control in a diverse population of children and youth with diabetes.

**Study design**—This was a cross-sectional analysis of data from a 6-center US study of diabetes in youth, including 3947 individuals with type 1 diabetes (T1D) and 552 with type 2 diabetes (T2D), using hemoglobin  $A_{1c}$  (Hb $A_{1c}$ ) levels to assess glycemic control.

**Results**—HbA<sub>1c</sub> levels reflecting poor glycemic control (HbA<sub>1c</sub> 9.5%) were found in 17% of youth with T1D and in 27% of those with T2D. African-American, American Indian, Hispanic, and Asian/Pacific Islander youth with T1D were significantly more likely to have higher HbA<sub>1c</sub> levels compared with non-Hispanic white youth (with respective rates for poor glycemic control of 36%, 52%, 27%, and 26% vs 12%). Similarly poor control in these 4 racial/ethnic groups was

Reprint requests: Ronny Bell, PhD, PI, SEARCH Coordinating Center, Department of Epidemiology and Prevention, Wake Forest University School of Medicine, Medical Center Boulevard, Winston-Salem, NC 27157-1063. rbell@wfubmc.edu.

<sup>\*</sup>A complete list of the members of the SEARCH for Diabetes in Youth Study Group is available at www.jpeds.com (Appendix 1). The SEARCH for Diabetes in Youth Study is indebted to the many youth and their families, as well as their health care providers, whose participation made this study possible.

Funding and conflict of interest information available at www.jpeds.com (Appendix 2).

found in youth with T2D. Longer duration of diabetes was significantly asso\*ciated with poorer glycemic control in youth with T1D and T2D.

**Conclusions**—The high percentage of US youth with  $HbA_{1c}$  levels above the target value and with poor glycemic control indicates an urgent need for effective treatment strategies to improve metabolic status in youth with diabetes.

Intensive glycemic control prevents the development or delays the progression of microvascular complications of diabetes in adults with type 1 diabetes (T1D) and type 2 diabetes  $(T2D)^{1,2}$  and in adolescents with T1D.<sup>3</sup> Lower HbA<sub>1c</sub> levels also reduce the risk of macrovascular disease in patients with T1D,<sup>4</sup> although recent results for patients with T2D are equivocal.<sup>5–7</sup>

In the Swedish Childhood Diabetes Registry (adjusted to the Diabetes Control and Complications Trial standard), for more than 3000 patients age < 20 years, the average hemoglobin  $A_{1c}$  (Hb $A_{1c}$ ) value was < 8% in 35% of the patients and > 9% in 29%.<sup>8</sup> Correlates of relatively high Hb $A_{1c}$  included female sex, older age, longer duration of diabetes, and high insulin dose. This type of descriptive data from large, unselected cohorts of youth with diabetes is critical to identifying groups of patients who may benefit from targeted interventions to improve metabolic control and thus reduce risk for long-term complications of diabetes. The SEARCH for Diabetes in Youth Study is a large observational study of childhood diabetes that includes a highly diverse population of youth with T1D and T2D. In the present work, we investigated the prevalence and correlates of good, intermediate, and poor glycemic control, measured using Hb $A_{1c}$ .

#### Methods

The SEARCH for Diabetes in Youth Study is ongoing at 6 study centers in the United States, with the goal of describing the epidemiology of childhood diabetes according to race/ ethnicity, age, sex, and diabetes type. The study design has been published previously.<sup>9</sup> It involves identifying existing (prevalent) cases of non-gestational diabetes in patients under age 20 years in 2001 and newly diagnosed (incident) cases in subsequent calendar years, with the goal of complete case ascertainment in each population under surveillance by the 6 study centers. The institutional review boards for all 6 sites approved the study protocol, and all activities are HIPAA-compliant. Prevalence for 2001<sup>10</sup> and incidence rates for 2002–2003 have been published,<sup>11</sup> with estimated case ascertainment completeness exceeding 90%.

The present analysis includes the 2001 prevalent and 2002–2005 incident study cohort participants with a clinical diagnosis of either T1D or T2D, as determined by each participant's health care provider. Data were collected for these cohorts between 2002 and 2007. Concerted efforts were made to contact each of the 11 179 patients with diabetes identified by the study in 2001–2005 whose diabetes was not secondary to other conditions to solicit their participation in an initial survey to collect information on age at diagnosis and race/ethnicity. The individuals who completed this survey were then asked to participate in an in-person research clinic visit that included blood sampling for HbA<sub>1c</sub> and other measures, a brief physical examination (including height and weight measurements), and an

interview dealing with socio-demographic factors and health issues. At the time of the study visit, informed consent was obtained from each participant age 18 or older and from the parent/guardian of any participant age 17 or younger. All measures were conducted by trained, certified staff in accordance with standardized study protocols (available at www.searchfordiabetes.org). HbA<sub>1c</sub> was measured in whole blood with an automated non-porous ion-exchange high-performance liquid chromatography system (model G-7; Tosoh Bioscience, Montgomeryville, Pennsylvania). This method has demonstrated to be linear from a total area of 500 to > 4500, indicating that the results are accurate within a large range of number of red cells. If the total area is < 500, then results are not reported; if the total area is > 4500, then the analysis is repeated after sample dilution. The intrassay coefficient of variation is 0.047%, the interassay coefficient of variation is 0.070%, and the normal reference range values are 4.2% to 5.8%.<sup>9</sup> Ultimately, 5299 (47%) of the 2001–2005 cases attended the research clinic visit. Not all of these individuals agreed to the blood draw; a total of 4499 individuals (3947 with T1D and 552 with T2D) had complete data and

### contributed data to the analysis. GAD65 was positive in 53.6% of the youth with T1D and in 18.9% of those with T2D, similar to previously reported data from SEARCH.<sup>11</sup>

#### Variable Definition

American Diabetes Association (ADA) target values for HbA<sub>1c</sub> in relation to age are as follows: 7.5% to 8.5% at age < 6 years, < 8.0% at age 6 to 12 years, < 7.5% at age 13 to 18 years, and < 7.0% at age 19+ years.<sup>12,13</sup> Individuals who met the ADA target (or for age < 6 years, who had an HbA<sub>1c</sub> < 8.5%) were classified as "good" control; those with HbA<sub>1c</sub> 9.5% regardless of age were classified as "poor" control, and those with HbA<sub>1c</sub> values between the definition of "good" and "poor" control were classified as "intermediate" control. HbA<sub>1c</sub> also was used in its continuous (uncategorized) form for statistical testing.

Height and weight measurements were used to calculate body mass index (BMI; in kg/m<sup>2</sup>). Age- and sex-specific BMI *z*-scores were derived from the Centers for Disease Control and Prevention (CDC) national standards, and the following weight status categories were assigned: "underweight or normal weight" for individuals < 85th percentile, "overweight" for those in the 85th to 95th percentiles, and "obese" for those > 95th percentile.<sup>14</sup> Self-reported race and ethnicity were collected using 2000 US Census questions. All participants who reported "Hispanic" ethnicity were categorized as "Hispanic," regardless of race. Among non-Hispanics, those who reported more than one race were placed into a single race category using the plurality approach of the National Center for Health Statistics.<sup>15</sup>

Parental education was defined as the highest educational level attained by either parent. Insurance source was categorized as "none," "private" (including private only or private plus something else), "Medicaid or Medicare," and "other." The latter category included Indian Health Service, military, school-based, and any other source not in combination with either private insurance or Medicaid/Medicare.

#### **Statistical Analysis**

All analyses were conducted using a single measure of  $HbA_{1c}$ , collected at the study examination. Key characteristics that can possibly affect glycemic control, including

underlying etiology, age at diagnosis, and diabetes treatment regimen, differ dramatically between youth with T1D and those with T2D. Because the intent of the present study was not to compare and contrast characteristics between diabetes types, but rather to describe glycemic control in youth with diabetes, analyses were conducted after stratification by diabetes type.

Subject characteristics were described using counts and percentages, stratified by diabetes type. Univariate associations between the subject characteristics and glycemic control (HbA<sub>1c</sub>) were tested for statistical significance using 1-way analysis of variance stratified by diabetes type. The *P* values for these associations were based on HbA<sub>1c</sub> as a continuous outcome, because this approach has greater statistical power.

Separate multivariate linear regression models (stratified by diabetes type) were used to evaluate the associations between each of the following subject characteristics and  $HbA_{1c}$  after adjusting for all other listed characteristics: age at study examination, duration of diabetes, weight status, family structure, diabetes care provider, race/ethnicity, sex, household income, parental education, and insurance source.

Although many statistical tests were conducted, because the present work was intended to be descriptive and hypothesis-generating in nature, the traditional P value < .05 (2-tailed test) was considered statistically significant. No formal correction was made for multiple tests. All statistical analyses were conducted using SAS for Windows version 9.1 (SAS Institute, Cary, North Carolina).

#### Results

The overall mean HbA<sub>1c</sub> value was  $8.18\% \pm 1.59\%$  for youth with T1D and  $7.99\% \pm 2.51\%$  for youth with T2D. Overall, 17% of the youth with T1D and 27% of those with T2D had poor glycemic control (ie, HbA<sub>1c</sub> 9.5%) (Table I; available at www.jpeds.com). For both T1D and T2D, the percentage of youth above the age-specific target HbA<sub>1c</sub> was higher with increasing age at the time of the SEARCH examination. In those age 19+ years, 29% of those with T1D and 47% of those with T2D exhibited poor glycemic control.

In univariate comparisons for T1D, glycemic control (ie, HbA<sub>1c</sub>) was significantly associated with all of the characteristics except weight status (Table I). After adjustment for age at the study examination, duration of diabetes, weight status, family structure, diabetes care provider, race/ethnicity, sex, household income, parental education, and insurance source, most patterns of association and statistical significance remained as observed in the unadjusted models. Multivariate results are presented in Table II (available at www.jpeds.com). Exceptions were weight status, which became statistically significant, and insurance source and household income, which were no longer statistically significant in the multivariate regression model. The statistically significant correlates of poorer glycemic control in the multivariate model for T1D were younger age, longer diabetes duration, weight <85th percentile (vs being obese), living in a single-parent household or other household structure (vs living in a 2-parent household), type of diabetes care provider (adult

endocrinologist or none vs pediatric endocrinologist), race/ethnicity other than non-Hispanic white, being female, and lower parental education (Table II).

Among participants with T2D, the descriptive univariate findings (Table I) revealed worse glycemic control in those with older age, longer duration of diabetes, normal-weight/ underweight or overweight status, "other" household structure (vs 2-parent or single-parent household), race/ethnicity other than non-Hispanic white, parental education less than high school or bachelor's degree or more, and no or "other" health insurance, whereas those cared for by a pediatric endocrinologist had better glycemic control. In the multivariate results (Table II), patterns of association were generally similar, although only duration of diabetes and parental education were statistically significant.

#### Discussion

A high proportion of children and youth with diabetes in this study exhibited poor  $HbA_{1c}$  values. This finding is particularly disturbing given that almost all of the youth were insured and all were motivated to volunteer for research.

Our finding of poor glycemic control in youth with T1D is similar to published data from other countries.<sup>8,16</sup> In these countries, there were center variations in glycemic control that were not explained by demographic or clinical factors. It has been suggested that a more detailed exploration of then implementation of treatment regimens may be informative.<sup>16</sup> In a separate report from SEARCH, youth with T1D who used insulin pumps had lower HbA<sub>1c</sub> values and fewer acute complications compared with those on other insulin regimens.<sup>17</sup>

The pattern of worsening glycemic control with increasing duration of T1D, independent of many other potential correlates, likely is due in part to progressive loss of beta cell function.<sup>18</sup> The difficulty of maintaining motivation for the intensive daily diabetes care patterns and lifestyle changes required to achieve glycemic targets likely is a contributing factor. Worse glycemic control in normal or underweight youth with T1D compared with their obese counterparts has not been reported previously, and reasons for this finding are unknown. The poorer residual beta cell function in youth with T1D with lower BMI<sup>18</sup> may play a role. Females had significantly worse glycemic control than males (Table II), although from a clinical perspective, the difference in HbA<sub>1c</sub> between the sexes was small (0.10).

Studies of children and youth with diabetes generally report a constellation of related sociodemographic factors associated with glycemic control, including race/ethnicity, socioeconomic status, parental education, parental involvement in diabetes management, and family dynamics. In the present study, African-American, Hispanic, American Indian, and Asian/Pacific Islander youth all had poorer glycemic control than non-Hispanic whites even after adjustment for all other variables studied. Comparing African-American and Caucasian children with T1D, Chalew et al<sup>19</sup> also reported higher mean HbA<sub>1c</sub> levels in the African-American children independent of sex, insurance status, BMI, and number of clinic visits. In contrast, however, Gallegos-Macias et al<sup>20</sup> reported that the higher HbA<sub>1c</sub> values seen in Hispanic youth with T1D compared with non-Hispanic white youth with T1D were

accounted for by lower socioeconomic status irrespective of race/ethnicity. Indeed, in the present study, lower parental education level and living in a single-parent or other family structure were associated independently with worse glycemic control. These factors may act either directly or indirectly through their influence on adherence to recommended selfcare.<sup>19–24</sup>

In univariate analyses, uninsured youth with T1D had poorer glycemic control, although after adjustment for other characteristics, this association was no longer statistically significant, perhaps due to the small number of youth with diabetes who were without insurance. Despite the fact that virtually all patients with T1D were insured, lower income was marginally associated with worse HbA1c values, even after adjusting for parental education, race/ethnicity, and clinical characteristics. Unmeasured financial impacts of insurance benefit structure—uncovered out-of-pocket expenses, copayments, and lost wages -affect families with various incomes differently, which might explain our observation. Low and modest income also may affect the ability of youth and their families to manage diabetes for reasons other than the monetary costs of health care, possibly including impaired access to diabetes care providers. Indeed, receiving diabetes care from a pediatric endocrinologist or diabetologist was associated independently with better glycemic control. Economic and other barriers to care will be the topic of further study in the SEARCH cohort. Improved understanding of the social mediators of the association between sociodemographic characteristics and glycemic control could assist in the development of tailored treatment strategies that might make it possible for patients and families to better adhere to diabetes care regimens and to attain their target HbA1c goals more easily.

Patterns of the correlates of glycemic control were generally similar for youth with T2D and those with T1D. But in the multivariate analyses, only duration of diabetes and attained parental education were statistically significant, likely due, at least in part, to lower statistical power given the substantially smaller number of subjects with T2D (n = 552) compared with those with T1D (n = 3947). Rothman et al<sup>25</sup> reported that among adolescents with T2D, after adjustment for a several demographic and clinical factors, HbA1c values were higher in their non-Caucasian subjects than in their Caucasian subjects. The present analysis adjusted for 2 variables that may partly account for that racial/ethnic disparity in glycemic control that were not assessed in the study of Rothman et al<sup>25</sup>—parental education and family structure-which may explain why in the present analysis, race/ethnicity was not statistically significantly associated with glycemic control. Results for educational attainment were somewhat unexpected, in that after adjustment for other factors, youth with T2D whose highest parental educational level was less than high school appeared to have comparable glycemic control with those with at least one parent with a bachelor's degree or higher, and better glycemic control was observed for those with intermediate levels of parental education. It may be that small sample size, particularly for the highest education grouping, generated a spurious result. As for youth with T1D, further study of the sociodemographic factors that affect the glycemic control of youth with T2D is needed. In addition, it is possible that the underlying genetic and biological factors that contribute variously to the etiology of diabetes (whether T1D or T2D) also may affect the relative ease or difficulty of meeting HbA1c targets. Such speculation should be the target of future investigations.

Limitations of the present study include the selective nature of the SEARCH centers and nonparticipation in the study visit at which blood is drawn, which might limit the generalizability of our results. The potential impact of nonresponse<sup>9</sup> on the present analysis was evaluated using routine clinical laboratory HbA1c test results from one of the study centers with institutional review board–approved access to clinical results for all patients who would be eligible to participate in the SEARCH study protocol. At this center, 1209 of the 1390 youth with diabetes in the 2001 and 2002 SEARCH study cohorts (87%) underwent HbA1c testing as part of their clinical care. For the youth in the 2001 prevalent cohort, the mean HbA1c was significantly lower in those who attended the study visit compared with those who did not ( $8.9\% \pm 1.9\%$  vs  $9.5\% \pm 2.4\%$ ), although for youth in the 2002 incident cohort, the results did not differ ( $9.5\% \pm 2.5\%$  vs  $9.4\% \pm 2.4\%$ ). Thus, our findings may underestimate the proportion of youth with diabetes in poor glycemic control.

Our HbA<sub>1c</sub> analyzer is linear over the large red blood cell range (total area, 500 to > 4500); however, we cannot exclude the possibility that a few individuals had aberrant results due to glycation, hemoglobin variants, and/or red cell life span.<sup>26</sup> We did not measure hematocrit or look for hemglobin variants; however, in persons with impaired glucose tolerance, adjustment for hematocrit and other factors likely to affect glycemiv control do not account for race/ethnic differences in HbA<sub>1c</sub>.<sup>27</sup> The most common cause of an aberrant value (albeit still rare) would be sickle cell anemia in the African-American subgroup,<sup>28</sup> in which red cell survival is decreased, resulting in lower HbA<sub>1c</sub> values than would be expected in relation to average blood glucose concentrations.

Strengths of the present study include its sample size, although despite the inclusion of > 500 youth with T2D, the limited variability in some characteristics may have limited the study's statistical power to detect potential clinically important differences in this T2D subgroup. Additional study strengths include the ethnic and geographic diversity and the use of a single laboratory to measure HbA<sub>1c</sub>.

Our data highlight the need for strategies to improve glycemic control in youth with diabetes. Technologies for managing diabetes continue to evolve.<sup>29,30</sup> Continuous glucose monitoring can now be used in conjunction with insulin pumps and traditional glucose monitoring by fingerstick to optimize glycemic control throughout the day. As with any diabetes care regimen, the financial and social burden on the patient and his or her family to maintain good metabolic control is substantial. Results from the Diabetes Control and Complications Trial indicate that glycemic control in participants on the intensive treatment arm was improved significantly by the consistent use of a nutrition plan relative to insulin dose.<sup>31</sup> Physical activity is another key determinant of glucose excursions and must be considered in optimal insulin dosing.<sup>32</sup>

Particular challenges arise when attempting to develop comprehensive diabetes management strategies that adequately address the complexity of diabetes care during adolescence, as physiological, emotional, and social development is unfolding. Recent successful interventions designed specifically for adolescents have used motivational interviewing<sup>33</sup> and behavioral family systems therapy for diabetes.<sup>34</sup> Further research is urgently needed to establish interventions that meld efficacious technology with effective behavioral and social

approaches to improve glycemic control for the highly diverse group of youth living with diabetes.

#### Glossary

BMI	Body mass index
HbA1c	Glycated hemoglobin
T1D	Type 1 diabetes
T2D	Type 2 diabetes

#### References

- Diabetes Control and Complications Trial Research Group. . The effect of intensive treatment of diabetes on the development and progression of long-term complications in insulin-dependent diabetes mellitus. N Engl J Med. 1993; 329:977–86. [PubMed: 8366922]
- UK Prospective Diabetes Study (UKPDS) Group. Intensive blood-glucose control with sulphonylureas or insulin compared with conventional treatment and risk of complications in patients with type 2 diabetes (UKPDS 33). Lancet. 1998; 352:837–53. [PubMed: 9742976]
- Diabetes Control and Complications Trial Research Group. . Effect of intensive diabetes treatment on the development and progression of long-term complications in adolescents with insulindependent diabetes mellitus: Diabetes Control and Complications Trial. J Pediatr. 1994; 125:177– 88. [PubMed: 8040759]
- Nathan DM, Cleary PA, Backlund JY, Genuth SM, Lachin JM, Orchard TJ, et al. Intensive diabetes treatment and cardiovascular disease in patients with type 1 diabetes. N Engl J Med. 2005; 353:2643–53. [PubMed: 16371630]
- Patel A, MacMahon S, Chalmers J, Neal B, Billot L, Woodward M, et al. Intensive blood glucose control and vascular outcomes in patients with type 2 diabetes. N Engl J Med. 2008; 358:2560–72. [PubMed: 18539916]
- Gerstein HC, Miller ME, Byington RP, Goff DC Jr, Bigger JT, Buse JB, et al. Effects of intensive glucose lowering in type 2 diabetes. N Engl J Med. 2008; 358:2545–59. [PubMed: 18539917]
- Holman RR, Paul SK, Bethel MA, Matthews DR, Neil HA. Ten-year follow-up of intensive glucose control in type 2 diabetes. N Engl J Med. 2008; 359:1577–89. [PubMed: 18784090]
- Hanberger L, Samuelsson U, Lindblad B, Ludvigsson J. A1C in children and adolescents with diabetes in relation to certain clinical parameters: the Swedish Childhood Diabetes Registry SWEDIABKIDS. Diabetes Care. 2008; 31:927–9. [PubMed: 18235047]
- 9. SEARCH for Diabetes in Youth: a multicenter study of the prevalence, incidence and classification of diabetes mellitus in youth. Control Clin Trials. 2004; 25:458–71. [PubMed: 15465616]
- Liese AD, D'Agostino RB Jr, Hamman RF, Kilgo PD, Lawrence JM, Liu LL, et al. The burden of diabetes mellitus among US youth: prevalence estimates from the SEARCH for Diabetes in Youth Study. Pediatrics. 2006; 118:1510–8. [PubMed: 17015542]
- 11. Dabelea D, Bell RA, D'Agostino RB Jr, Imperatore G, Johansen JM, Linder B, et al. Incidence of diabetes in youth in the United States. JAMA. 2007; 297:2716–24. [PubMed: 17595272]
- Silverstein J, Klingensmith G, Copeland K, Plotnick L, Kaufman F, Laffel L, et al. Care of children and adolescents with type 1 diabetes: a statement of the American Diabetes Association. Diabetes Care. 2005; 28:186–212. [PubMed: 15616254]
- American Diabetes Association. Type 2 diabetes in children and adolescents. Diabetes Care. 2000; 23:381–9. [PubMed: 10868870]
- 14. Centers for Disease Control and Prevention. CDC Growth Charts: United States, 2000. Atlanta, GA: Centers for Disease Control and Prevention; 2000.

- Ingram DD, Parker JD, Schenker N, Weed JA, Hamilton B, Arias E, et al. United States Census 2000 population with bridged race categories. Vital Health Stat 2. 2003; 135:1–55. [PubMed: 14556588]
- 16. de Beaufort CE, Swift PG, Skinner CT, Aanstoot HJ, Aman J, Cameron F, et al. Continuing stability of center differences in pediatric diabetes care: do advances in diabetes treatment improve outcome? The Hvidoere Study Group on Childhood Diabetes. Diabetes Care. 2007; 30:2245–50. [PubMed: 17540955]
- 17. Paris CA, Imperatore G, Klingensmith G, Petitti D, Rodriguez B, Ruggiero A, et al. Predictors of insulin regimens and impact on outcomes in youth with type 1 diabetes: the SEARCH for Diabetes in Youth Study. J Pediatr. 2009; 155:182–9.
- Scholin A, Bjorklund L, Borg H, Arnqvist H, Bjork E, Blohme G, et al. Islet antibodies and remaining beta-cell function 8 years after diagnosis of diabetes in young adults: a prospective follow-up of the nationwide Diabetes Incidence Study in Sweden 5. J Intern Med. 2004; 255:384– 91. [PubMed: 14871463]
- Chalew SA, Gomez R, Butler A, Hempe J, Compton T, Mercante D, et al. Predictors of glycemic control in children with type 1 diabetes: the importance of race. J Diabetes Complications. 2000; 14:71–7. [PubMed: 10959068]
- 20. Gallegos-Macias AR, Macias SR, Kaufman E, Skipper B, Kalishman N. Relationship between glycemic control, ethnicity and socioeconomic status in Hispanic and white non-Hispanic youths with type 1 diabetes mellitus. Pediatr Diabetes. 2003; 4:19–23. [PubMed: 14655519]
- 21. Danne T, Mortensen HB, Hougaard P, Lynggaard H, Aanstoot HJ, Chiarelli F, et al. Persistent differences among centers over 3 years in glycemic control and hypoglycemia in a study of 3,805 children and adolescents with type 1 diabetes from the Hvidore Study Group. Diabetes Care. 2001; 24:1342–7. [PubMed: 11473067]
- 22. Moreland EC, Tovar A, Zuehlke JB, Butler DA, Milaszewski K, Laffel LM. The impact of physiological, therapeutic and psychosocial variables on glycemic control in youth with type 1 diabetes mellitus. J Pediatr Endocrinol Metab. 2004; 17:1533–44. [PubMed: 15570991]
- 23. Forsander GA, Sundelin J, Persson B. Influence of the initial management regimen and family social situation on glycemic control and medical care in children with type I diabetes mellitus. Acta Paediatr. 2000; 89:1462–8. [PubMed: 11195237]
- Anderson B, Ho J, Brackett J, Finkelstein D, Laffel L. Parental involvement in diabetes management tasks: relationships to blood glucose monitoring adherence and metabolic control in young adolescents with insulin-dependent diabetes mellitus. J Pediatr. 1997; 130:257–65. [PubMed: 9042129]
- 25. Rothman RL, Mulvaney S, Elasy TA, VanderWoude A, Gebretsadik T, Shintani A, et al. Selfmanagement behaviors, racial disparities, and glycemic control among adolescents with type 2 diabetes. Pediatrics. 2008; 121:e912–9. [PubMed: 18381520]
- Cohen RM, Franco RS, Khera PK, Smith EP, Lindsell CJ, Ciraolo PJ, et al. Red cell life span heterogeneity in hematologically normal people is sufficient to alter HbA1c. Blood. 2008; 112:4284–91. [PubMed: 18694998]
- 27. Herman WH, Ma Y, Uwaifo G, Haffner S, Kahn SE, Horton ES, et al. Differences in A1C by race and ethnicity among patients with impaired glucose tolerance in the Diabetes Prevention Program. Diabetes Care. 2007; 30:2453–7. [PubMed: 17536077]
- Obesity and cardiovascular disease risk factors in black and white girls: the NHLBI Growth and Health Study. Am J Public Health. 1992; 82:1613–20. [PubMed: 1456335]
- 29. Steck AK, Klingensmith GJ, Fiallo-Scharer R. Recent advances in insulin treatment of children. Pediatr Diabetes. 2007; 8(Suppl 6):49–56. [PubMed: 17727385]
- Tamborlane WV, Beck RW, Bode BW, Buckingham B, Chase HP, Clemons R, et al. The Juvenile Diabetes Research Foundation Continuous Glucose Monitoring Study Group. Continuous glucose monitoring and intensive treatment of type 1 diabetes New England. Journal of Medicine. 2008; 359:1464–76.
- Delahanty LM, Halford BN. The role of diet behaviors in achieving improved glycemic control in intensively treated patients in the Diabetes Control and Complications Trial. Diabetes Care. 1993; 16:1453–8. [PubMed: 8299434]

- Tansey MJ, Tsalikian E, Beck RW, Mauras N, Buckingham BA, Weinzimer SA, et al. The effects of aerobic exercise on glucose and counterregulatory hormone concentrations in children with type 1 diabetes. Diabetes Care. 2006; 29:20–5. [PubMed: 16373890]
- 33. Channon SJ, Huws-Thomas MV, Rollnick S, Hood K, Cannings-John RL, Rogers C, et al. A multicenter randomized controlled trial of motivational interviewing in teenagers with diabetes. Diabetes Care. 2007; 30:1390–5. [PubMed: 17351283]
- Wysocki T, Harris MA, Buckloh LM, Mertlich D, Lochrie AS, Mauras N, et al. Randomized trial of behavioral family systems therapy for diabetes: maintenance of effects on diabetes outcomes in adolescents. Diabetes Care. 2007; 30:555–60. [PubMed: 17327320]

#### Appendix 1

Members of the Search for Diabetes in Youth Study Group. California: Jean M. Lawrence, ScD, MPH, MSSA, Ann K. Kershnar, MD, Kristi Reynolds, PhD, MPH, and Marlene Y. Gonzalez, MPH, for Kaiser Permanente Southern California; David J. Pettitt, MD, for the Sansum Diabetes Research Institute; and Diana B. Petitti, MD, MPH, for the University of Southern California

**Colorado:** Dana Dabelea, MD, PhD, Richard F. Hamman, MD, DrPH, and Lisa Testaverde, MS, for the Department of Preventive Medicine and Biometrics, University of Colorado Denver; Georgeanna J. Klingensmith, MD and Marian J. Rewers, MD, PhD, for the Barbara Davis Center for Childhood Diabetes; Stephen Daniels, MD, PhD, Department of Pediatrics and Children's Hospital, University of Colorado Denver School of Medicine; Clifford A. Bloch, MD, for Pediatric Endocrine Associates; Jonathan Krakoff, MD and Peter H. Bennett, MD, FRCP, for the NIDDK Pima Indian Study; Joquetta A. DeGroat, BA, for the Navajo Area Indian Health Prevention Program; and Teresa Coons, PhD, for St. Mary's Hospital Grand Junction

**Hawaii:** Beatriz L. Rodriguez, MD, PhD, Beth Waitzfelder, PhD, Wilfred Fujimoto, MD, J. David Curb, MD, Fiona Kennedy, RN, Greg Uramoto, MD, Sorrell Waxman, MD, Teresa Hillier, MD, and Richard Chung, MD, for the Pacific Health Research Institute

**Ohio:** Lawrence M. Dolan, MD, Michael Seid, PhD, Nancy Crimmins, MD, and Debra A. Standiford, MSN, CNP for the Cincinnati Children's Hospital Medical Center

**South Carolina:** Elizabeth J. Mayer-Davis, PhD and Joan Thomas MS, RD for the University of North Carolina Chapel Hill; Angela D. Liese, PhD, MPH, Robert McKeown, PhD, Robert R. Moran, PhD, Deborah Truell, RN, CDE, Gladys Gaillard-McBride, RN, CFNP, Deborah Lawler, MT (ASCP), and Malaka Jackson, MD for the University of South Carolina; Lynne Hartel, MA, Yaw Appiagyei-Dankah, MD, and Lyndon Key, MD, for the Medical University of South Carolina; Sheree Mejia, RN, James Amrhein, MD, and Kent Reifschneider, MD, for Greenville Hospital Systems; Pam Clark, MD for McLeod Pediatric Subspecialists; Mark Parker, MD for Pediatric Endocrinology & Diabetes Specialists; and I. David Schwartz, MD for Pediatric Endocrinology at the Medical College of Georgia

**Washington:** Catherine Pihoker, MD, Lisa Gilliam, MD, PhD, Irl Hirsch, MD, Lenna L. Liu, MD, MPH, Carolyn Paris, MD, MPH, and Dimitri Christakis, MD, MPH for the University of Washington; Beth Loots, MPH, MSW, Joyce Yi, PhD, Stacey Bryant, RN,

Michelle Sadler-Greever, RN, CDE, Rebecca O'Connor, RN, Ellen Braun-Kelly, BS, Amber Sexton, BS, and Corinne Shubin, BA for the Seattle Children's Hospital and Regional Medical Center; and Carla Greenbaum, MD for the Benaroya Research Institute

**Centers for Disease Control and Prevention:** Giuseppina Imperatore, MD, PhD, Desmond E. Williams, MD, PhD, Michael M. Engelgau, MD, Henry S. Kahn, MD, K. M. Venkat Narayan, MD, MPH, and Bernice Moore, MBA

National Institute of Diabetes and Digestive and Kidney Diseases, National Institutes of Health: Barbara Linder, MD, PhD

**Central Laboratory (University of Washington):** Santica M. Marcovina, PhD, ScD, Vinod P. Gaur, PhD, and Kathy Gadbois

**Coordinating Center (Wake Forest University School of Medicine):** Ronny Bell, PhD, MS, Ralph D'Agostino, Jr, PhD, Douglas Case, PhD, Timothy Morgan, PhD, Michelle J. Naughton, PhD, Susan Vestal, BS, Gena Hargis, MPH, Andrea Anderson, MS, Cralen Davis, MS, Jeanette Andrews, MS, and Jennifer Beyer, MS

#### Appendix 2

The SEARCH for Diabetes in Youth Study is funded by the Centers for Disease Control and Prevention (CDC) (PA number 00097 and DP-05-069) and supported by the National Institute of Diabetes and Digestive and Kidney Diseases (NIDDK). Site contract numbers are as follows: Kaiser Permanente Southern California, U01 DP000246; University of Colorado Health Sciences Center, U01 DP000247; Pacific Health Research Institute, U01 DP000245; Children's Hospital Medical Center (Cincinnati), U01 DP000248; University of North Carolina at Chapel Hill, U01 DP000254; University of Washington School of Medicine, U01 DP000244; Wake Forest University School of Medicine, U01 DP000250. The authors acknowledge the involvement of general clinical research centers at the following institutions in the SEARCH for Diabetes in Youth Study: Medical University of South Carolina (Grant M01 RR01070), Cincinnati Children's Hospital (Grant M01 RR08084), Children's Hospital and Regional Medical Center and the University of Washington School of Medicine (Grants M01RR00037 and M01RR001271), and Colorado Pediatric General Clinical Research Center (Grant M01 RR00069). The contents of this article are solely the responsibility of the authors and do not necessarily represent the official views of the CDC or the NIDDK. The authors declare no conflicts of interest.

Author Manuscript

## Table I

Percentage of individuals with T1D or T2D with good, intermediate, or poor glycemic control according to clinical and demographic characteristics: SEARCH for Diabetes in Youth, prevalent 2001 and incident 2002-2005 study participants

			<b>T1D</b> ( $n = 3947$ )					T2D $(n = 552)$		
		Ċ	cemic control, %	*,0			Gly	cemic control, %	*.0	
Characteristic	Z	Good	Intermediate	Poor	$P^{\dagger}$	u	Good	Intermediate	Poor	$P^{\dagger}$
All	3947	44.4	38.8	16.8		552	53.8	19.6	26.6	
Age at diagnosis, years					<.0001					.1529
0–5	1185	44.8	40.1	15.1		0		I		
6-12	2065	43.6	38.9	17.5		182	58.2	17.6	24.2	
13–18	693	45.7	36.7	17.6		360	51.9	20.8	27.2	
19+	4	75.0	0.0	25.0		10	40.0	10.0	50.0	
Age at examination, years					<.0001					<.0001
0–5	402	6.99	25.1	8.0		0		I	I	
6-12	1748	54.1	34.7	11.3		LL	72.7	11.7	15.6	
13–18	1499	32.4	44.4	23.3		369	57.5	19.5	23.0	
19+	298	17.8	53.7	28.5		106	27.4	25.5	47.2	
Diabetes duration, months					<.0001					<.0001
< 12	1167	69.4	23.1	7.5		183	71.0	16.4	12.6	
12–23	827	47.8	37.1	15.1		164	56.7	20.1	23.2	
24-47	581	38.9	43.0	18.1		113	44.3	19.5	36.3	
48+	1369	23.5	51.3	25.3		92	26.1	25.0	48.9	
Weight status <sup>‡</sup>					.2712					.0002
Underweight or normal weight (<85th percentile)	2654	44.4	39.0	16.6		67	43.3	11.9	44.8	
Overweight (85th to 94th percentile)	808	41.8	40.5	17.7		69	46.4	21.7	31.9	
Obese (> 95th percentile)	485	48.7	35.1	16.3		416	56.7	20.4	22.8	
Family structure					<.0001					.0088
Two-parent household	2623	50.1	37.7	12.2		224	52.2	20.1	27.7	
Single-parent household	1126	33.9	41.5	24.6		238	59.7	18.9	21.4	
Other household structure	177	26.0	38.4	35.6		57	43.9	21.1	35.1	
Diabetes care provider					<.0001					<.0001

			<b>T1D</b> $(n = 3947)$					T2D (n = 552)		
		Gly	cemic control, <sup>9</sup>	*•%			Glyc	cemic control, %	*,0	:
Characteristic	Z	Good	Intermediate	Poor	$P^{\dagger}$	u	Good	Intermediate	Poor	$P^{\dagger}$
Pediatric endocrinologist or diabetologist	2995	45.5	39.0	15.4		309	65.4	17.2	17.5	
Adult endocrinologist	151	18.5	45.7	35.8		31	41.9	12.9	45.2	
General pediatrician, family physician, or general internist	146	28.8	41.1	30.1		91	33.0	25.3	41.8	
Nurse practitioner or physician assistant	552	50.7	34.6	14.7		40	55.0	20.0	25.0	
Other/don't know	75	36.0	42.7	21.3		37	40.5	29.7	29.7	
None, no medical care	8	25.0	37.5	37.5		12	16.7	16.7	66.7	
Race/ethnicity					<.0001					<.0001
Non-Hispanic white	2983	46.9	40.8	12.3		107	71.0	16.8	12.2	
African American	355	34.7	29.9	35.5		175	58.9	18.9	22.3	
Hispanic	440	39.1	33.6	27.3		117	50.4	22.2	27.4	
Asian/Pacific Islander	127	37.0	37.0	26.0		4	47.7	15.9	36.4	
American Indian	23	17.4	30.4	52.2		105	34.3	21.9	43.8	
Sex					.0040					.1610
Female	1961	42.9	39.2	17.9		350	50.3	21.4	28.3	
Male	1986	45.8	38.4	15.8		202	59.9	16.3	23.8	
Household income					<.0001					.6613
<\$25 K	495	35.6	34.3	30.1		201	57.2	20.4	22.4	
\$25–49 K	848	40.6	39.2	20.3		129	53.5	24.8	21.7	
\$50–74 K	793	46.0	40.9	13.1		56	64.3	8.9	26.8	
\$75 + K	1500	50.9	39.5	9.6		48	66.7	10.4	22.9	
Parental education					<.0001					.0160
Less than high school	160	34.4	28.1	37.5		87	46.0	18.4	35.6	
High school graduate or GED	618	35.8	38.0	26.2		172	61.1	19.2	19.8	
Some college (but less than bachelor's degree)	1296	41.3	40.1	18.6		164	55.5	22.0	22.6	
Bachelor's degree or more	1843	50.5	39.1	10.4		89	50.6	16.9	32.6	
Insurance					<.0001					.0013
None	59	28.8	30.5	40.7		21	28.6	28.6	42.9	
Private	3127	46.5	39.8	13.7		266	57.9	16.9	25.2	
Medicaid/Medicare	666	36.3	35.1	28.5		191	56.5	22.5	20.9	

Author Manuscript

			T1D $(n = 3947)$					T2D (n = 552)		
		Ē	ycemic control, 9	*•			Gly	cemic control, %	*.0	
Characteristic	Z	Good	Intermediate	Poor	$P^{\dagger}$	u	Good	Intermediate	Poor	$P^{\dagger}$
Other§	69	42.0	36.2	21.7		40	35.0	20.0	45.0	

Glycemic control defined as "good" used age-specific HbA1c target values as follows: <6 years, <8.5%; 6 to 12 years, <8.0%; 13 to 18 years, <7.5%; 19+ years, <7.0%. "Poor" glycemic control was defined as HbA1c 9.5%. "Intermediate" glycemic control was defined as values between "good" and "poor."

 $\dot{\tau}$  values are based on analysis of variance, treating HbA1c as a continuous outcome and testing the association with each covariate individually.

 $\sharp$  Weight status defined based on CDC guidelines using age- and sex-specific BMI percentiles (Ref).

 $^{\$}$ Includes Indian Health Service, military, school-based, and other (when these are not in combination with either private insurance or Medicaid/Medicare).

Associations of HbA<sub>1c</sub> with clinical and demographic characteristics (adjusted): SEARCH for Diabetes in Youth, prevalent 2001 and incident 2002–2005 study participants

		T1D (n = 3606)			<b>T2D</b> ( <b>n</b> = 421)	
Characteristic	Estimate	95% CI	$P^*$	Estimate	95% CI	$P^*$
Age at examination, years			<.0001			.1470
0–5	Ref	I				
6–12	-0.33	-0.49 to -0.16	<.0001	0.66	-1.60 to 0.28	.1679
13–18	-0.15	-0.32 to 0.03	.0956	-0.75	-1.50 to $0.00$	.0515
19+	-0.56	-0.84 to -0.28	<.0001	Ref		
Diabetes duration, months			<.0001			.0058
< 12	Ref	I		Ref		
12–23	0.67	0.54 to 0.80	<.0001	0.41	-0.14 to 0.96	.1417
24-47	0.89	0.74 to 1.03	<.0001	0.72	0.08 to 1.36	.0286
48+	1.18	1.05 to 1.30	<.0001	1.38	0.60 to 2.17	.0006
Weight status $^{\dagger}$			.0018			.2378
Underweight or normal weight (< 85th percentile)	0.19	0.05 to 0.34	9600.	0.57	-0.15 to 1.29	.1231
Overweight (85th to 94th percentile)	0.02	-0.15 to 0.19	.8255	0.33	-0.36 to 1.02	.3447
Obese (> 95th percentile)	Ref	I		Ref		
Family structure			.0003			.1866
Two-parent household	Ref	I		Ref		
Single-parent household	0.19	0.08 to 0.30	.0011	-0.42	-0.91 to 0.07	.0896
Other household structure	0.39	0.14 to 0.65	.0024	0.01	-0.80 to 0.83	.9746
Diabetes care provider			.0049			.1218
Pediatric endocrinologist or diabetologist	Ref	I		Ref		
Adult endocrinologist	0.41	0.10 to 0.72	0600.	0.34	-0.79 to 1.47	.5517
General pediatrician or family physician or general internist	0.21	-0.05 to 0.47	.1069	0.49	-0.32 to 1.30	.2346
Nurse practitioner or physician assistant	-0.06	-0.19 to 0.07	.3831	0.78	-0.01 to 1.56	.0530
Other/don't know	-0.07	-0.42 to 0.29	.7130	-0.12	-1.19 to 0.95	.8221
None, no medical care	1.48	0.41 to 2.54	.0068	1.96	0.18 to 3.75	.0310
Race/ethnicity			<.0001			.1322

		T1D (n = 3606)			(174 - II) (171	
Characteristic	Estimate	95% CI	$P^*$	Estimate	95% CI	$P^*$
Non-Hispanic white	Ref		I	Ref		
African American	0.69	0.52 to 0.87	<.0001	0.52	-0.09 to 1.14	.0959
Hispanic	0.25	0.09 to 0.40	.0025	0.47	-0.19 to 1.13	.1652
Asian/Pacific Islander	0.41	0.14 to 0.67	.0027	06.0	0.03 to 1.77	.0418
American Indian	1.02	0.34 to 1.71	.0034	1.16	0.10 to 2.21	.0327
Sex			.0277			.7680
Female	0.10	0.01 to 0.20	.0277	0.07	-0.38 to 0.52	.7680
Male	Ref			Ref		
Household income			.0597			.4847
<\$25 K	Ref			Ref	I	I
\$25–49 K	-0.13	-0.31 to 0.04	.1323	-0.04	-0.61 to 0.52	.8813
\$50-74 K	-0.22	-0.42 to -0.02	.0328	0.17	-0.61 to 0.96	.6622
\$75 + K	-0.27	-0.47 to -0.07	.0081	-0.53	-1.42 to 0.36	.2434
Parental education			<.0001			.0447
Less than high school	0.40	0.12 to 0.68	.0052	-0.07	-0.90 to 0.77	.8694
High school graduate or GED	0.35	0.19 to 0.50	<.0001	-0.75	-1.43 to -0.07	.0300
Some college (but less than bachelor's degree)	0.26	0.15 to 0.38	<.0001	-0.64	-1.30 to 0.01	.0554
Bachelor's degree or more	Ref			Ref	Ι	I
Insurance			.2499			.8844
None	0.38	-0.04 to 0.79	.0763	0.20	-1.12 to 1.53	.7617
Private	Ref			Ref	Ι	I
Medicaid/Medicare	0.03	-0.13 to 0.20	.6946	-0.18	-0.75 to 0.39	.5347
Other <sup>‡</sup>	-0.17	-0.54 to 0.20	.3720	-0.25	-1.42 to 0.92	.6760

J Pediatr. Author manuscript; available in PMC 2015 December 23.

Petitti et al.

 $t^{\pm}$ Includes Indian Health Service, military, school-based, and other (when these are not in combination with either private insurance or Medicaid/Medicare).

 $^{\dagger}$  Weight status defined based on CDC guidelines using age- and sex-specific BMI percentiles (Ref).

Author Manuscript