Bucknell University

Bucknell Digital Commons

Faculty Journal Articles

Faculty Scholarship

2018

Associations of Four Community Factors with Longitudinal Change in Hemoglobin A1c Levels in Patients with Type 2 **Diabetes**

Elizabeth Durden Bucknell University, tdurden@bucknell.edu

Follow this and additional works at: https://digitalcommons.bucknell.edu/fac_journ



Part of the Medicine and Health Commons

Recommended Citation

Durden, Elizabeth. "Associations of Four Community Factors with Longitudinal Change in Hemoglobin A1c Levels in Patients with Type 2 Diabetes." Diabetes Care (2018): 461-468.

This Article is brought to you for free and open access by the Faculty Scholarship at Bucknell Digital Commons. It has been accepted for inclusion in Faculty Journal Articles by an authorized administrator of Bucknell Digital Commons. For more information, please contact dcadmin@bucknell.edu.



Associations of Four Community Factors With Longitudinal Change in Hemoglobin A_{1c} Levels in Patients With Type 2 Diabetes

Diabetes Care 2018;41:461-468 | https://doi.org/10.2337/dc17-1200

Annemarie G. Hirsch,^{1,2} T. Elizabeth Durden,³ Cara Nordberg,⁴ Andrea Berger,⁴ and Brian S. Schwartz^{1,2}

OBJECTIVE

To evaluate associations of community factors with glycated hemoglobin (HbA_{1c}).

RESEARCH DESIGN AND METHODS

We identified patients with type 2 diabetes who had an $HbA_{1c} \ge 7.5\%$ (58 mmol/mol) and subsequent HbA_{1c} testing within 90–270 days. We used mixed-effect models to assess whether treatment intensification (TI) and community domains (community socioeconomic deprivation [CSD], food availability, fitness assets, and utilitarian physical activity favorability [quartiled]) were associated with HbA_{1c} change over 6 and 24 months, controlling for demographics, HbA_{1c} , BMI, and time with evidence of type 2 diabetes. We evaluated whether community domains modified associations of TI with HbA_{1c} change using cross product terms.

RESULTS

There were 15,308 patients with 69,818 elevated HbA_{1c} measures. The average reduction in HbA_{1c} over 6 months was 0.07% less in townships with a high level of CSD (third quartile versus the first). Reductions were 0.10% greater for HbA_{1c} in townships with the best food availability (versus worst). HbA_{1c} reductions were 0.17–0.19% greater in census tracts in the second and third quartiles of utilitarian physical activity favorability versus the first. The association of TI with 6-month HbA_{1c} change was weaker in townships and boroughs with the worst CSD (versus best) and in boroughs with the best fitness assets (versus worst). The association of TI with 24-month HbA_{1c} change was weaker in census tracts with the worst CSD (versus third quartile) and strongest in census tracts most favorable for utilitarian physical activity (versus worst).

CONCLUSIONS

Community domains were associated with HbA_{1c} change and blunted TI effectiveness.

Control of glycated hemoglobin (HbA $_{1c}$) is a primary therapeutic target in type 2 diabetes (1,2). While HbA $_{1c}$ control among persons with diabetes has improved in the U.S. over the past two decades, >40% of persons with type 2 diabetes have HbA $_{1c}$ values >7.0% (53 mmol/mol) (3), the target recommended by treatment guidelines (4). Prior work has examined associations between clinic-, physician-, and patient-level factors and HbA $_{1c}$ control (5). However, very little is known about whether where a person lives is associated with HbA $_{1c}$ control.

Patients with type 2 diabetes are particularly vulnerable to the influence of communitylevel factors because of the importance of self-management that takes place largely Corresponding author: Annemarie G. Hirsch, aghirsch@geisinger.edu.

Received 15 June 2017 and accepted 20 November 2017.

© 2017 by the American Diabetes Association. Readers may use this article as long as the work is properly cited, the use is educational and not for profit, and the work is not altered. More information is available at http://www.diabetesjournals.org/content/license.

¹Department of Epidemiology and Health Services Research, Geisinger Health System, Danville,

²Department of Environmental Health and Engineering, Johns Hopkins Bloomberg School of Public Health, Baltimore, MD

³Department of Sociology and Anthropology, Bucknell University. Lewisbura. PA

⁴Department of Biomedical and Translational Informatics, Geisinger Health System, Danville, PA

outside of the clinical setting. Prior ecological studies have reported that the community poverty rate, unemployment rate, population density, and other characteristics have been found to explain variation in type 2 diabetes prevalence across counties (6,7); however no study, to our knowledge, has examined how these factors impact HbA_{1c} control at the individual level. A recent study reported that supermarket presence in communities was associated with HbA_{1c} trajectories among patients with type 2 diabetes (8). However, to date no one has evaluated how physical activity opportunities and food availability beyond supermarkets (e.g., fast food outlets) impact HbA_{1c} levels over time among individuals with type 2 diabetes.

We conducted a study of >15,000 patients with type 2 diabetes living in 734 communities in Pennsylvania to assess whether the food environment, physical activity environment, or community socioeconomic deprivation (CSD) were associated with HbA_{1c} levels over time. We also examined whether these factors modified associations between medical care and changes in HbA_{1c}. This question is fundamentally important to identifying barriers and facilitators of successful patient outcomes.

RESEARCH DESIGN AND METHODS

We conducted a retrospective cohort study of primary care patients from the Geisinger with type 2 diabetes. We used data from 2001-2015 from the Geisinger electronic health record (EHR) and community measures to evaluate 1) association of type 2 diabetes treatment intensification (TI) with change in HbA_{1c} levels, 2) associations of community factors with change in HbA_{1c} levels, and 3) whether associations of TI with change in HbA_{1c} levels were modified by key community factors. Community features under study included CSD, food availability, fitness and recreational assets, and utilitarian physical activity favorability.

Study Population and Subject Selection

Geisinger is an integrated health system that serves >500.000 primary care patients across Pennsylvania and New Jersey. The primary care population is representative of the general population in the region (8). We classified patients as having type 2 diabetes using a validated EHR-based

algorithm (9): at least two clinical encounters associated with a type 2 diabetes diagnosis (ICD-9: 250.00-250.90, with 5th digit either 0 or 2) on two separate dates, a type 2 diabetes medication order and at least two elevated laboratory tests (random glucose ≥200 mg/dL, fasting glucose \geq 126 mg/dL, or HbA_{1c} \geq 6.5% [53 mmol/mol]), a type 2 diabetes encounter diagnosis and a type 2 diabetes medication order, or a type 2 diabetes encounter diagnosis and an elevated HbA_{1c} value.

Of 249,123 primary care patients between 18 and 75 years of age, 35,260 patients met the criteria for type 2 diabetes. Of these patients, 2,220 were excluded if they had the following evidence of type 1 diabetes including: more than one type 1 diabetes diagnosis (ICD-9: 250.01-250.91, with 5th digit either 1 or 3) associated with a clinical encounter, a type 1 encounter diagnosis after the last recorded type 2 encounter diagnosis, or type 1 and type 2 diagnoses recorded on the same day. We then removed patients with <365 days between first and last encounters in the EHR (n = 4,942), patients without outpatient visits (n = 23), and patients without an HbA_{1c} value \geq 7.5% (58 mmol/mol) (n = 10,172). Among the remaining 17,903 patients, there were 125,335 elevated HbA_{1c} values; for these, there was not a follow-up for 22,009 within 90-270 days, insulin was prescribed for an additional 32,356 prior to the HbA_{1c} measure, and 1,152 occurred within 30 days of each other. After removal of these HbA_{1c} measures, the final analytical data set consisted of 15,308 patients with a total of 69,818 HbA_{1c} measures.

Study Variables

HbA_{1c} Outcome

The dependent variable for this analysis was change in HbA_{1c} level over time. We first identified elevated HbA_{1c} levels for patients as values ≥7.5% (58 mmol/mol). We then searched for subsequent HbA_{1c} values measured between 90 and 270 days after the elevated value. We calculated the change in HbA_{1c} as the difference between the elevated measure and the subsequent measure closest to 6 months after baseline (follow-up HbA_{1c} – baseline HbA_{1c} = HbA_{1c} change); a negative value for change in HbA_{1c} indicated an improvement in HbA_{1c} levels. Six-month HbA_{1c} values could also qualify as baseline HbA_{1c} values if they were still elevated.

TI

TI measures monitor how providers respond to information about control of risk factors. For type 2 diabetes, we defined TI as a change in medication regimen in response to an elevated HbA_{1c} level meeting one of the following criteria: 1) an increase in the number of medication classes ordered, 2) an increased dose of a medication, or 3) a switch to a medication in a different therapeutic class (10,11). We excluded HbA_{1c} values from analysis if a patient had an insulin order prior to the date of the elevated HbA_{1c} level, as day-to-day adjustments in insulin cannot be identified in the EHR.

To operationalize the TI measure, we determined baseline medication use by diabetes medication orders prior to the date of the elevated HbA_{1c}. Next, we classified an occurrence of any of the following as TI if the change occurred on or within 30 days after the date of the baseline HbA_{1c}: 1) initiation of insulin, 2) addition of a diabetes medication in a subclass not on the patient's baseline list, or 3) increase of dose (dose/pill \times pills/day) of a baseline medication.

Medication dose was only available in a free-text field in the EHR. Natural language processing was applied to identify common dose entries (i.e., entries found in at least 35 medication orders). These common entries were then manually classified as TI or no TI. Entries found for <35 medication orders were classified as "unknown TI" as manual review of the large number of unique free-text entries was not feasible.

Community Measures

Using previously described methods, we geocoded study subjects to their residential address in the EHR (12). Each home address was assigned to 1 of 734 communities using two sets of U.S. census-defined boundaries: minor civil divisions and census tracts. Minor civil divisions include townships, which range from rural to suburban; boroughs, which are generally walkable small- to medium-sized towns; and cities, which are large and heterogeneous geographies. We used a mixed definition in identifying communities, assigning patients to townships, boroughs, and census tracts in cities, as previously reported (8,12).

We conducted a confirmatory factor analysis (CFA) to develop a theory-based, formal model for measuring latent constructs that characterized obesity-related environments. Based on prior literature, care.diabetesjournals.org Hirsch and Associates 463

we selected an initial pool of >150 candidate indicators (e.g., density, diversity, and accessibility measures) from archival data from the Geisinger service area. We estimated the CFA model using progressive model refinement to achieve an acceptable fit. Once the optimal model was identified, we generated factor scores for each of four constructs: 1) CSD (proportion of population in poverty, unemployed, or on public assistance, with less than a high school education, and not in labor force), 2) food availability (food service establishments per population; convenience stores, including in gasoline service stations, per area; grocery stores per area; snack food stores per area; other food retail stores per area; bars and taverns per population; fast food chain restaurants per area; and diversity of types of food establishments), 3) fitness and recreational assets (indoor fitness clubs per area, indoor/outdoor recreational clubs per area, diversity of recreational and fitness establishments, and count of outdoor public parks), and 4) utilitarian physical activity favorability (households per area, vehicle miles traveled per population, average block size [square miles], and street connectivity [percentage of road intersections that are connected]). Density measures for food and fitness assets were computed as the number of establishments per square mile. Diversity measures were based on 13 categories of food establishments (e.g., fast food chain, bars, taverns) and 9 categories of physical activity establishments (e.g., bowling, golf courses). For each community, the number of categories with one or more facilities was counted.

CSD was derived from U.S. Census data; food and physical activity establishment data were based on InfoUSA and Dun & Bradstreet data; and utilitarian physical activity favorability data were based on data from the Pennsylvania Department of Transportation. While the original CFA used data from these sources from the year 2000, for this study we used 2010 data (most recently available data), z-transforming and summing the same indicators to create updated factor scores. Each of the four pairs of 2000 and 2010 factor scores were moderately to highly correlated (Spearman values ranged from 0.6458 to 0.8075).

Statistical Analysis

The goals of the analysis were to identify the main effect of TI on HbA_{1c} change,

identify the main effects of community domains on HbA_{1c} change, and evaluate moderation of TI main effect by community domains. First, we created a linear regression model to evaluate the association of TI on change in HbA_{1c} , controlling for baseline HbA_{1c} (continuous, centered, linear, and quadratic); sex; age (continuous, centered, linear, and quadratic); race/ethnicity (non-Hispanic white vs. all others); BMI (continuous); duration of type 2 diabetes in the EHR, defined as the time between the date of first type 2 diabetes diagnosis code, diabetes medication, or elevated HbA_{1c} and the date of the elevated baseline HbA_{1c} (years as continuous, centered, linear, and quadratic); and Medical Assistance (MA) (ever vs. never received) status for health insurance as a surrogate for family socioeconomic status (SES) (13,14). Comorbid diseases (e.g., hypertension, coronary artery disease, and fatty liver disease) were evaluated for confounding and were not retained in the final models. We included a random intercept for person to account for multiple HbA_{1c} change measures per person. Next, we created four mixedeffect linear regression models to evaluate associations of each of the four community domains (in quartiles) with HbA_{1c} change.

These models also included a random intercept for person and controlled for the same covariates described above, as well as TI status. Finally, we added a cross product term between TI and each community factor to each model to assess effect modification. For examination of the durability of the associations observed at 6 months, the analysis was repeated for HbA_{1c} change over 24 months.

We stratified all analysis by community type (i.e., township, borough, and city census tract), since distributions of environmental measures differed by geography, leading to regression extrapolation. We did not include a random intercept for community, as the intraclass correlation coefficients for the models were very low (0.005–0.006). All analysis was performed with SAS software, version 9.2, of the SAS Institute

RESULTS

Description of Study Population

There were 15,308 eligible patients who had at least one elevated HbA_{1c} and a follow-up HbA_{1c} value within the specified time period (Table 1). The mean age of patients was 57.8 years of age, 45.8% of patients were female, and the majority of patients were white. MA was used by

Table 1—Characteristics of patients with type 2 diabetes with elevated HbA_{1c} and a follow-up HbA_{1c} within 90–270 days and characteristics of HbA_{1c} values

Characteristics	Patients (n = 15,308) ^a
Age, years	57.8 ± 11.9
Female, n (%)	7,006 (45.8)
Race/ethnicity, <i>n</i> (%) Non-Hispanic white All others	14,761 (96.4) 517 (3.4)
MA, n (%)	1,964 (12.8)
Elevated HbA_{1c} with follow-up value in specific window, number per person	3.0 (1.0–6.0)
Community type, <i>n</i> (%) Township Borough Census tract in city	9,063 (59.2) 4,342 (28.4) 1,590 (10.4)
Elevated ${\rm HbA_{1c}}$ values (n = 69,818) Initial elevated ${\rm HbA_{1c}}$ value, % Duration between elevated ${\rm HbA_{1c}}$ and follow-up value in	8.2 (7.7–9.1)
specific time window, days	174.60 ± 42.95
Change in HbA _{1c} in specified time window	-0.6 ± 1.5
TI, n (%)	24.206 (24.0)
Yes No	24,396 (34.9) 38,274 (54.8)
Unknown ^b	7,148 (10.2)

Data are mean \pm SD or median (interquartile range) unless otherwise indicated. ^aTotals may not add to 15,308 patients owing to missing/unknown values. ^bMedication regimens (e.g., dose and schedule) are entered as free-text in the EHR. We applied natural language processing to identify and classify TI status for commonly used text patterns. Unknown patterns included single-use test strings that could not be used to determine TI status without manual chart review.

12.8% of patients at some point while under care by Geisinger. The median number of eligible elevated ${\rm HbA_{1c}}$ measures per person was three for a total of 69,818 elevated measures. The mean time between the ${\rm HbA_{1c}}$ measures in each pair (i.e., elevated and follow-up) was 174.6 days. TI occurred for slightly more than one-third of elevated ${\rm HbA_{1c}}$ values.

Description of Communities

Patients resided in 734 communities (Table 2), with the majority in townships (9,063 patients in 439 townships), followed by boroughs (4,342 patients in 205 boroughs) and city census tracts (1,590 patients in 90 census tracts). Community domains varied within and across community types. CSD was generally highest in census tracts and lowest in townships. Similarly, food availability, fitness and recreational assets, and utilitarian physical activity favorability scores were generally highest in census tracts and lowest in townships.

Multilevel Regression Modeling of HbA_{1c} Change

Initial elevated HbA_{1c} values had a median of 8.2% (66 mmol/mol) followed by an absolute average decline of 0.6%. In our tabulated model results, coefficients

indicate the absolute difference in HbA_{1c} change between the variable level (e.g., TI) and the reference group (no TI), such that negative coefficients represent a larger decline in HbA_{1c} over time than the reference group and positive coefficients represent a smaller decline in HbA_{1c} over time than the reference group. In each community type, on average, TI (versus no TI) was associated with a larger adjusted absolute decline in HbA_{1c} over both 6 and 24 months, though the magnitude of the decline was smaller at 24 months (Table 3). Unknown TI (versus not) was associated with larger adjusted absolute decline in HbA_{1c} but only over 6 months.

The four community factors were then added to the TI models one at a time. Adding community factors to the TI models did not alter TI associations. In townships, CSD was associated with absolute change in $\rm HbA_{1c}$ in townships. On average, $\rm HbA_{1c}$ reductions over time were 0.07% smaller for $\rm HbA_{1c}$ measures of patients living in communities with bad CSD (third quartile) compared with the best CSD (first quartile) (P < 0.05). CSD was not associated with $\rm HbA_{1c}$ change in boroughs and cities. Adding MA to the models did not substantially change CSD

associations. This association was not significant at 24 months.

Community food availability was associated with change in HbA_{1c} in townships only, with average absolute declines that were 0.1% larger for patients with the best versus the worst food availability. This same finding was observed at 24 months. Utilitarian physical activity was associated with change in HbA_{1c} in census tracts only, with average absolute declines over 6 months that were 0.17-0.19% larger for patients in communities at the second or third quartile (versus first). At 24 months, the average absolute decline was 0.25% larger for patients in census tracts at the third quartile (versus first.) The was no evidence that fitness assets were associated with change in HbA_{1c} at 6 months, but at 24 months townships with more physical activity opportunities had HbA_{1c} reductions that were 0.11% smaller than in townships with fewer opportunities (second quartile vs. first).

Effect Modification by Community Factor

There was evidence that TI was associated with smaller improvements in HbA_{1c} over 6 months in boroughs and townships with the worst CSD (fourth vs. first quartile)

Factors and indicators	Township ($n = 439$)	Borough ($n = 205$)	Census tract $(n = 90)$
CSD Population in poverty (%, z score) Civilian unemployment rate (%, z score) Public assistance (%, z score) <high (%,="" education="" force="" in="" labor="" not="" school="" score)="" score)<="" td="" z=""><td>-1.03 (-6.27, 8.82)</td><td>-0.05 (-5.26, 12.09)</td><td>3.21 (-2.91, 18.24)</td></high>	-1.03 (-6.27, 8.82)	-0.05 (-5.26, 12.09)	3.21 (-2.91, 18.24)
	-0.44 (-1.44, 4.91)	0.08 (-1.49, 4.44)	0.87 (-0.96, 6.48)
	-0.27 (-1.92, 3.40)	-0.02 (-1.92, 4.48)	0.29 (-1.63, 3.49)
	-0.41 (-1.01, 3.71)	-0.08 (-1.01, 5.59)	0.73 (-1.01, 6.65)
	-0.20 (-2.13, 5.36)	-0.21 (-1.85, 3.11)	0.59 (-1.25, 3.87)
	-0.22 (-2.03, 5.81)	-0.06 (-1.89, 4.86)	0.04 (-1.08, 4.24)
Community food availability Food service per 1,000 persons Convenience/gas station per square mile Grocery store area per square mile Snack food stores per square mile Other food retail per square mile Bar and tavern population per square mile Fast food chain restaurant per square mile Diversity of types of food establishments	-2.59 (-4.71, 17.40) 0.44 (0.00, 11.82) 0.00 (0.00, 2.03) 0.00 (0.00, 2.37) 0.00 (0.00, 3.38) 0.00 (0.00, 4.06) 0.00 (0.00, 5.46) 0.00 (0.00, 2.03) 2.00 (0.00, 13.00)	0.64 (-4.81, 13.69) 2.11 (0.00, 6.71) 0.66 (0.00, 11.40) 0.53 (0.00, 10.18) 0.00 (0.00, 11.27) 0.40 (0.00, 9.50) 0.00 (0.00, 2.47) 0.00 (0.00, 3.03) 4.00 (0.00, 11.00)	4.85 (-3.92, 56.17) 2.29 (0.00, 17.03) 2.45 (0.00, 22.43) 2.21 (0.00, 50.72) 0.95 (0.00, 24.99) 2.70 (0.00, 24.99) 0.56 (0.00, 4.39) 0.00 (0.00, 6.09) 6.00 (1.00, 11.00)
Fitness and recreational assets Indoor fitness clubs per square mile Outdoor recreational clubs per square mile Indoor recreational clubs per square mile Diversity of fitness/recreational places	-2.05 (-3.2, 15.55)	-0.55 (-3.26, 14.92)	3.76 (-2.65, 31.98)
	0.00 (0.00, 1.35)	0.00 (0.00, 14.37)	0.67 (0.00, 17.95)
	0.00 (0.00, 1.01)	0.00 (0.00, 4.53)	0.00 (0.00, 11.66)
	0.00 (0.00, 0.63)	0.00 (0.00, 7.18)	0.00 (0.00, 13.88)
	2.00 (0.00, 9.00)	2.00 (0.00, 9.00)	3.00 (0.00, 8.00)
Count of outdoor public parks/recreational spaces Utilitarian physical activity favorability Households per square mile	0.00 (0.00, 5.00)	0.00 (0.00, 2.00)	0.00 (0.00, 2.00)
	-0.46 (-16.11, 2.98)	0.85 (-2.23, 3.59)	1.95 (-1.68, 8.71)
	27.33 (0.39, 860.92)	644.55 (27.60, 4,134.04)	2,188.52 (188.17, 9,159.5)
Vehicle miles traveled per person per square mile	23.64 (3.63, 433.27)	9.16 (0.10, 105.49)	6.13 (0.67, 63.77)
Average block size (square miles)	0.41 (0.02, 3.56)	0.02 (0.00, 0.73)	0.01 (0.00, 0.11)
Street connectivity: number of 3+ intersections/m ²	0.09 (0.00, 0.23)	0.05 (0.00, 0.19)	0.02 (0.00, 0.11)

Data are median (range) unless stated otherwise. aU.S. census data, InfoUSA and Dun & Bradstreet, and the Pennsylvania Department of Transportation.

care.diabetesjournals.org Hirsch and Associates 46

Table 3—Associations of TI and community factors for change in HbA_{1c} in 6-month^a window after an elevated HbA_{1c} measure, by community type^b

continuity type			
	Township, estimate (95% CI)	Borough, estimate (95% CI)	Census tract, estimate (95% CI)
TI in model without any community factors			
Yes vs. no	-0.34 (-0.36, -0.31) ^c	$-0.34 (-0.36, -0.29)^{c}$	$-0.42 (-0.48, -0.35)^{c}$
Unknown vs. no	-0.08 (-0.11, -0.04) ^c	−0.11 (−0.16, −0.05) ^c	-0.11 (-0.21, -0.011) ^c
Main effect of community factors: in models			
one at a time (quartile by place)			
CSD			
Quartile 2	0.04 (-0.02, 0.09)	0.01 (-0.08, 0.10)	0.03 (-0.13, 0.19)
Quartile 3	0.07 (0.01, 0.13) ^c	0.05 (-0.03, 0.14)	0.07 (-0.08, 0.23)
Quartile 4	0.06 (-0.002, 0.11)	0.04 (-0.04, 0.13)	0.06 (-0.11, 0.23)
Utilitarian PA favorability			
Quartile 2	0.04 (-0.03, 0.10)	0.01 (-0.10, 0.11)	-0.19 (-0.35, -0.03) ^c
Quartile 3	0.03 (-0.04, 0.09)	-0.01 (-0.11, 0.09)	-0.17 (-0.34, -0.004) ^c
Quartile 4	0.02 (-0.04, 0.07)	0.04 (-0.06, 0.14)	0.03 (-0.13, 0.20)
Community food availability			
Quartile 2	-0.0004 (-0.07, 0.07)	-0.03 (-0.15, 0.08)	-0.04 (-0.10, 0.21)
Quartile 3	-0.05 (-0.12, 0.02)	0.02 (-0.08, 0.13)	0.05 (-0.10, 0.21)
Quartile 4	$-0.10 (-0.16, -0.03)^{c}$	-0.03 (-0.13, 0.07)	0.12 (-0.02, 0.26)
Fitness and recreational assets			
Quartile 2	0.06 (-0.02, 0.13)	-0.08 (-0.21, 0.06)	-0.05 (-0.21, 0.10)
Quartile 3	-0.03 (-0.10, 0.04)	-0.04 (-0.16, 0.09)	-0.002 (-0.16, 0.15)
Quartile 4	-0.05 (-0.11, 0.02)	-0.08 (-0.20, 0.04)	0.10 (-0.06, 0.26)
Effect modification on TI by community factors:			
in models one at a time (quartile by place) ^d			
CSD quartile 2 $ imes$ TI yes	0.02 (-0.04, 0.09)	0.03 (-0.08, 0.14)	-0.07 (-0.27, 0.14)
CSD quartile 3 $ imes$ TI yes	0.04 (-0.02, 0.11)	-0.09 (-0.02, 0.19)	-0.03 (-0.22, 0.17)
CSD quartile 4 $ imes$ TI yes	0.11 (0.05, 0.18) ^c	0.11 (0.007, 0.21) ^c	-0.06 (-0.28, 0.16)
Fitness quartile 2 $ imes$ TI yes	0.05 (-0.04, 0.13)	0.18 (0.01, 0.35) ^c	0.05 (-0.15, 0.24)
Fitness quartile 3 $ imes$ TI yes	-0.02 (-0.06, 0.10)	0.11 (-0.05, 0.27)	-0.08 (-0.27, 0.11)
Fitness quartile 4 $ imes$ TI yes	-0.009 (-0.08, 0.06)	0.16 (0.007, 0.31) ^c	0.05 (-0.15, 0.24)

PA, physical activity. a HbA $_{1c}$ value closest to 6 months after baseline HbA $_{1c}$. b Mixed-effect linear regression models with a random intercept for patient, adjusted for race/ethnicity and MA, age centered (linear and quadratic), baseline HbA $_{1c}$ (linear and quadratric), sex, BMI centered, and time from diabetes diagnosis to baseline HbA $_{1c}$ (linear and quadratic); change in HbA $_{1c}$ was calculated as follow-up HbA $_{1c}$ minus initial elevated HbA $_{1c}$. Reference group for community measures was quartile 1. Reference group for TI was "no TI." $^{c}P < 0.05$. d Cross product of community factors and TI added to linear regression models one at a time. Results presented for community factors with at least one significant (P < 0.05) community \times TI yes interaction term.

(Table 4). This same pattern was observed at 24 months but only in census tracts (third vs. first quartile) (Table 4). TI was also associated with smaller improvements in HbA_{1c} in boroughs with the highest level of fitness assets (fourth quartile vs. first quartile) (Table 3). Notably, TI was associated with greater improvements in HbA_{1c} over 24 months in census tracts that had the highest level of utilitarian physical activity opportunities (second and fourth quartile vs. first quartile) (Table 4).

CONCLUSIONS

This is the first study, to our knowledge, to evaluate the association between community characteristics and control of HbA_{1c} over time and how these characteristics modify HbA_{1c} response to care. We found that in certain community types, CSD, food availability, and utilitarian physical activity favorability were associated with HbA_{1c} trajectories. Moreover,

community factors blunted the associations of TI with ${\rm HbA_{1c}}$ levels. The results suggest that factors outside of the clinical setting may be important to ${\rm HbA_{1c}}$ control and point to opportunities for more targeted secondary prevention strategies that take community into account.

To date, most studies of community and type 2 diabetes have focused on risk of onset. Living in communities with greater resources to support physical activity and healthy diets has been associated with lower incidence of type 2 diabetes (15–17). Only two other studies to our knowledge have evaluated community characteristics and HbA_{1c} control over time in patients with type 2 diabetes. These studies found that loss of supermarket presence in communities and higher levels of chronic environmental contamination were associated with worse HbA_{1c} trajectories (7,8,18).

This study demonstrated that other aspects of the community, including

opportunities for utilitarian physical activity. the food environment, and CSD, were associated with HbA_{1c} trajectories, independent of treatment, in some community types. These associations are biologically plausible through a number of pathways. Communities with high levels of CSD, for example, create a context for higher psychosocial stress; stress has been linked to physiological changes associated with type 2 diabetes onset and higher HbA_{1c} (19-21). An obesity pathway may also explain our observed associations. Obesity is a strong risk factor for poor type 2 diabetes control, and weight loss in type 2 diabetes is associated with improvements in glycemic control (22,23). Community domains, including CSD and opportunities for physical activity, have been associated with obesity (13,24,25).

A decrease of 1% in HbA_{1c} has been associated with a 15–20% decrease in major cardiovascular events and a 37% decrease in microvascular complications

Table 4—Associations of TI and community factors for change in HbA_{1c} in 24-month^a window after an elevated HbA_{1c} measure, by community typeb

	Township, estimate (95% CI)	Borough, estimate (95% CI)	Census tract, estimate (95% CI)
TI in model without any community factors			
Yes vs. no	-0.13 (-0.15, -0.10) ^c	$-0.12 (-0.15, -0.08)^{c}$	-0.13 (-0.19, -0.07) ^c
Unknown vs. no	-0.01 (-0.05, 0.02)	-0.02 (-0.08, 0.03)	-0.05 (-0.15, 0.04)
Community factors: in models one at a time			
(quartile by place)			
CSD			
Quartile 2	0.03 (-0.04, 0.10)	0.002 (-0.11, 0.12)	-0.03 (-0.23, 0.18)
Quartile 3	-0.07 (-0.01, 0.14)	0.06 (-0.05, 0.17)	-0.01 (-0.21, 0.18)
Quartile 4	0.06 (-0.01, 0.13)	0.04 (-0.07, 0.15)	-00.18 (-0.40, 0.04)
Utilitarian PA favorability			
Quartile 2	0.001 (-0.07, 0.09)	-0.01 (-0.14, 0.12)	-0.20 (-0.41, 0.01)
Quartile 3	-0.03 (-0.11, 0.06)	0.01 (-0.12, 0.14)	-0.25 (-0.46, -0.03) ^c
Quartile 4	-0.02 (-0.09, 0.06)	0.07 (-0.06, 0.19)	-0.08 (-0.29, 0.13)
Community food availability			
Quartile 2	0.02 (-0.07, 0.10)	-0.04 (-0.19, 0.11)	-0.10 (-0.28, 0.07)
Quartile 3	-0.07 (-0.15, 0.02)	-0.02 (-0.16, 0.12)	0.02 (-0.18, 0.22)
Quartile 4	-0.10 (-0.19, -0.02) ^c	-0.04 (-0.17, 0.08)	-0.03 (-0.22, 0.15)
Fitness and recreational assets			
Quartile 2	0.11 (0.01, 0.20) ^c	-0.12 (-0.30, 0.05)	0.03 (-0.18, 0.23)
Quartile 3	0.03 (-0.06, 0.11)	-0.05 (-0.21, 0.10)	0.03 (-0.17, 0.23)
Quartile 4	-0.04 (-0.12, 0.04)	-0.09 (-0.25, 0.05)	0.04 (-0.16, 0.25)
Effect modification on TI by community factors:			
in models one at a time (quartile by place) ^d			
CSD quartile 2 $ imes$ TI yes	-0.002 (-0.07, 0.06)	-0.05 (-0.16, 0.06)	-0.16 (-0.36, 0.04)
CSD quartile 3 $ imes$ TI yes	-0.03 (-0.09, 0.04)	-0.03 (-0.14, 0.07)	-0.20 (-0.39, -0.005) ^c
CSD quartile 4 $ imes$ TI yes	0.009 (-0.05, 0.07)	-0.03 (-0.13, 0.07)	-0.10 (-0.31, 0.12)
Utilitarian PA quartile 2 $ imes$ TI yes	0.05 (-0.02, 0.12)	0.004 (-0.12, 0.13)	-0.30 (-0.50, -0.09) ^c
Utilitarian PA quartile 3 $ imes$ TI yes	-0.006 (-0.08, 0.07)	-0.01 (-0.14, 0.11)	-0.15 (-0.36, -0.06)
Utilitarian PA quartile 4 $ imes$ TI yes	0.02 (-0.05, 0.08)	-0.02 (-0.14, 0.10)	-0.29 (-0.49, -0.09) ^c
	1		

PA, physical activity. ${}^aHbA_{1c}$ value closest to 24 months after baseline HbA_{1c} . bMixed -effect linear regression models with a random intercept for patient, adjusted for race/ethnicity and MA, age centered (linear and quadratic), baseline HbA_{1c} (linear and quadratric), sex, BMI centered, and time from diabetes diagnosis to baseline HbA_{1c} (linear and quadratic); change in HbA_{1c} was calculated as follow-up HbA_{1c} minus initial elevated HbA_{1c}. Reference group for community measures was quartile 1. Reference group for TI was "no TI." $^cP < 0.05$. d Cross product of community factors and TI added to linear regression models one at a time. Results presented for community factors with at least one significant (P < 0.05) community \times TI yes interaction term.

(26). We observed that TI was associated with an absolute additional decline in HbA_{1c} of 0.35- 0.44% over 6 months, thus potentially resulting in up to an additional 8.8% decrease in major cardiovascular events. On average, the HbA_{1c} levels of patients living in census tracts favorable to utilitarian physical activity (quartile 3) dropped an average of 0.25% more than those in the communities least favorable to utilitarian physical activity over 24 months, a reduction that might be expected to produce a 4-5% reduction in cardiovascular disease and a 9% reduction in risk of microvascular complications.

The food and utilitarian physical activity environments, like TI, had durable associations with HbA_{1c} reductions. The magnitude of the association between TI and HbA_{1c} change was lower with longer durations between HbA_{1c} values; in contrast, the magnitude of associations with the food environment remained the same and strengthened for utilitarian

physical activity. The difference in durability may reflect the stability of environmental measures over these durations, providing for a more persistent role, while the impact of TI at a single clinical encounter, based on our findings, appears to weaken over time.

Consistent with prior studies, we demonstrated that TI was associated with improvements in HbA_{1c} over time (11,27). Sidorenkov et al. (28) reported that TI (versus none) was associated with a decline in HbA_{1c} of 0.21% after 21–120 days. Selby et al. (11) demonstrated that facilitylevel improvements in TI rates were associated with improvement in facilitylevel HbA_{1c} control rates. We also observed that unknown TI status resulted in greater HbA_{1c} reductions compared with patients known not to have had TI. We suspect that the unknown group included patients who actually received TI. To our knowledge, no study has evaluated how characteristics of where patients live modify the relation between TI and outcomes. We observed that CSD and opportunities for physical activity modified the effectiveness of diabetes care in some community types.

The mechanism for the observed effect modification is not clear. Worse CSD and neighborhood safety both have been associated with worse medication adherence rates and delays in filling prescriptions in type 2 diabetes (29,30). It may be that high CSD in townships and boroughs presents barriers to adherence, thus resulting in a weaker association between TI, as defined by medication orders, and HbA_{1c}. Regarding fitness assets, aerobic exercise has been shown to interact with a number of medications that are used to treat type 2 diabetes. influencing changes in insulin sensitivity, insulin secretion, or glycemic control (31).

While all four community domains that we studied were associated with HbA_{1c} or modified TI and HbA_{1c} associations, none care.diabetesjournals.org Hirsch and Associates 467

of our observed associations between community and HbA_{1c} existed in all three of our community types. There was very little overlap in the distribution of scores for the community domains across the three community types, with census tracts generally scoring the worst for CSD but best for food availability, fitness, and utilitarian physical activity—and townships the opposite extreme. It is possible that community factors have nonlinear relations with HbA_{1c} change, including threshold exposure effects, as has been observed for population density (32). Residents may only begin to use active transport (i.e., walking) above population densities of 3,500 persons per square mile. We evaluated associations by quartiling by community type; therefore, it may be that our highest quartiles of some community factors in some community types did not exceed thresholds needed to change behaviors. It is notable that there were instances when there was a statistically and clinically significant association at the 2nd or 3rd quartile of some community domains, but there was not a statistically significant association at the 4th quartile. It may be that there are unmeasured community factors that occur at the highest quartiles that counteract the benefits of HbA_{1c} reduction.

Less than 12% of HbA_{1c} variation among patients with diabetes can be attributed to physician- or clinical-level factors (5). Our findings demonstrated that while TI improved HbA_{1c} control, factors outside of the clinic were associated with HbA_{1c} control. Furthermore, we found that the association between TI and HbA_{1c} reduction was blunted by community characteristics. This finding suggests that community features may act like genetic differences; both community and genetic differences may modify the individual response to therapy, as has already been demonstrated in pharmacogenetic research (33). This finding is relevant to the growing precision medicine movement, an approach to medicine that selects treatments specific to the genes, environment, and lifestyle of each patient. Our results raise the possibility that medical treatments may need to be tailored to community characteristics to be effectively optimized (34).

The study used a unique combination of longitudinal treatment data, HbA_{1c} from laboratory measurements, and community-level variables on >15,000 individuals

across 734 communities. These data sources enabled us to evaluate associations between community factors and changes in ${\rm HbA_{1c}}$ levels while controlling for potential confounding variables. We also used multidimensional community measures from a CFA model, rather than single indicators in each domain, overcoming limitations of most prior research (35–38), which has generally relied on single indicators, ignoring the spatial co-occurrence of multiple features in each community domain.

The study had several limitations. First, we cannot conclude that our associations are generalizable outside of the region we studied. Second, self-selection may bias our results such that factors that motivate individuals to live in certain communities may be confounding our findings. However, we controlled for a number of variables known to be associated with diabetes status, including race/ethnicity, age, sex, and family SES. Third, we assessed community factors using geographic information systems but did not measure patient perceptions of their communities. Christine et al. (15) observed that geographic information system-based measures were less strongly associated with type 2 diabetes than measures of the perception of the availability of food and physical activity establishments. Finally, health care providers generally do not record health behaviors in the EHR, so we were unable to evaluate whether changes in health-related behaviors could have accounted for the

Despite improvements in type 2 diabetes care and treatment options, HbA_{1c} levels remain poorly controlled for many individuals with this disease. Our findings are a novel contribution to understanding correlates of type 2 diabetes control beyond clinical care. Our observed associations between community and HbA_{1c} levels, particularly that community factors may modify the influence of medical care, provide evidence to support the study of disease-management strategies that take community factors into consideration.

Funding. This study was funded by the Bucknell-Geisinger Research Initiative.

Duality of Interest. No potential conflicts of interest relevant to this article were reported. **Author Contributions**. A.G.H., T.E.D., and B.S.S. designed the study. A.G.H., A.B., and B.S.S. ob-

tained the study data. A.G.H., C.N., A.B., and B.S.S. analyzed the study data. A.G.H. wrote the manuscript. All authors reviewed and edited the manuscript. A.G.H. is the guarantor of this work and, as such, had full access to all the data in the study and takes responsibility for the integrity of the data and the accuracy of the data analysis.

References

- 1. American Diabetes Association. Glycemic targets. Sec. 6. In *Standards of Medical Care in Diabetes—2015*. Diabetes Care 2015;38(Suppl. 1):S33–S40
- 2. Palta P, Huang ES, Kalyani RR, Golden SH, Yeh HC. Hemoglobin A1c and mortality in older adults with and without diabetes: results from the National Health and Nutrition Examination Surveys (1988-2011). Diabetes Care 2017:40:453–460
- 3. Selvin E, Parrinello CM, Sacks DB, Coresh J. Trends in prevalence and control of diabetes in the United States, 1988-1994 and 1999-2010. Ann Intern Med 2014;160:517–525
- American Diabetes Association. Standards of Medical Care in Diabetes—2016: summary of revisions. Diabetes Care 2016;39(Suppl. 1):S4–S5
- O'Connor PJ, Rush WA, Davidson G, et al. Variation in quality of diabetes care at the levels of patient, physician, and clinic (Abstract). Prev Chronic Dis 2008:5:A15
- 6. Hipp JA, Chalise N. Spatial analysis and correlates of county-level diabetes prevalence, 2009-2010. Prev Chronic Dis 2015;12:E08
- 7. Zhang YT, Mujahid MS, Laraia BA, et al. Association between neighborhood supermarket presence and glycated hemoglobin levels among patients with type 2 diabetes mellitus. Am J Epidemiol 2017:185:1297–1303
- 8. Liu AY, Curriero FC, Glass TA, Stewart WF, Schwartz BS. The contextual influence of coal abandoned mine lands in communities and type 2 diabetes in Pennsylvania. Health Place 2013;22:115–122 9. Newton KM, Peissig PL, Kho AN, et al. Validation of electronic medical record-based phenotyping algorithms: results and lessons learned from the eMERGE network. J Am Med Inform Assoc 2013;20:e147–e154
- 10. Rose AJ, Berlowitz DR, Manze M, Orner MB, Kressin NR. Comparing methods of measuring treatment intensification in hypertension care. Circ Cardiovasc Qual Outcomes 2009;2:385–391
- 11. Selby JV, Uratsu CS, Fireman B, et al. Treatment intensification and risk factor control: toward more clinically relevant quality measures. Med Care 2009;47:395–402
- 12. Casey JA, Curriero FC, Cosgrove SE, Nachman KE, Schwartz BS. High-density livestock operations, crop field application of manure, and risk of community-associated methicillin-resistant Staphylococcus aureus infection in Pennsylvania. JAMA Intern Med 2013;173:1980–1990
- Nau C, Schwartz BS, Bandeen-Roche K, et al. Community socioeconomic deprivation and obesity trajectories in children using electronic health records. Obesity (Silver Spring) 2015;23:207–212
 Hirsch AG, Pollak J, Glass TA, et al. Early-life antibiotic use and subsequent diagnosis of food allergy and allergic diseases. Clin Exp Allergy 2017; 47:236–244
- 15. Christine PJ, Auchincloss AH, Bertoni AG, et al. Longitudinal associations between neighborhood physical and social environments and incident type 2 diabetes mellitus: the Multi-Ethnic

- Study of Atherosclerosis (MESA). JAMA Intern Med 2015:175:1311-1320
- 16. Creatore MI, Glazier RH, Moineddin R, et al. Association of neighborhood walkability with change in overweight, obesity, and diabetes. JAMA 2016;315:2211-2220
- 17. Astell-Burt T, Feng X, Kolt GS. Is neighborhood green space associated with a lower risk of type 2 diabetes? Evidence from 267,072 Australians. Diabetes Care 2014:37:197-201
- 18. Liu AY, Curriero FC, Glass TA, Stewart WF, Schwartz BS. Associations of the burden of coal abandoned mine lands with three dimensions of community context in Pennsylvania [article online], 2012. Available from https://www.hindawi.com/archive/ 2012/251201/. Accessed 8 December 2017
- 19. Gary-Webb TL, Suglia SF, Tehranifar P. Social epidemiology of diabetes and associated conditions. Curr Diab Rep 2013;13:850-859
- 20. Joseph JJ, Wang X, Spanakis E, et al. Diurnal salivary cortisol, glycemia and insulin resistance: the Multi-Ethnic Study of Atherosclerosis. Psychoneuroendocrinology 2015;62:327-335
- 21. Champaneri S. Xu X. Carnethon MR. et al. Diurnal salivary cortisol and urinary catecholamines are associated with diabetes mellitus: the Multi-Ethnic Study of Atherosclerosis. Metabolism 2012;61:986-995
- 22. Lyssenko V, Jonsson A, Almgren P, et al. Clinical risk factors, DNA variants, and the development of type 2 diabetes. N Engl J Med 2008;359:
- 23. Wing RR, Lang W, Wadden TA, et al.; Look AHEAD Research Group. Benefits of modest

- weight loss in improving cardiovascular risk factors in overweight and obese individuals with type 2 diabetes. Diabetes Care 2011;34:1481-
- 24. Nau C, Ellis H, Huang H, et al. Exploring the forest instead of the trees: an innovative method for defining obesogenic and obesoprotective environments. Health Place 2015;35:136-146
- 25. Schwartz BS, Stewart WF, Godby S, et al. Body mass index and the built and social environments in children and adolescents using electronic health records. Am J Prev Med 2011;41:e17-e28 26. Church TS. Blair SN. Cocreham S. et al. Effects of aerobic and resistance training on hemoglobin A1c levels in patients with type 2 diabetes: a randomized controlled trial, JAMA 2010:304:2253-
- 27. Berlowitz DR, Ash AS, Glickman M, et al. Developing a quality measure for clinical inertia in diabetes care. Health Serv Res 2005;40:1836-
- 28. Sidorenkov G, Voorham J, Haaijer-Ruskamp FM, de Zeeuw D, Denig P. Association between performance measures and glycemic control among patients with diabetes in a communitywide primary care cohort. Med Care 2013;51: 172-179
- 29. Billimek J, August KJ. Costs and beliefs: understanding individual- and neighborhood-level correlates of medication nonadherence among Mexican Americans with type 2 diabetes. Health Psvchol 2014:33:1602-1605
- 30. Billimek J, Sorkin DH. Self-reported neighborhood safety and nonadherence to treatment

- regimens among patients with type 2 diabetes. J Gen Intern Med 2012:27:292-296
- 31. Faerch K, Hulman A, Soloman TPJ. Heterogeneity of pre-diabetes and type 2 diabetes: implications for prediction, prevention, and treatment responsiveness. Curr Diabetes Rev 2016;12:30-41 32. Ross C, Dunning A. Land Use Transportation Interaction: An Examination of the 1995 NPTS Data. Washington, DC, Federal Highway Administration, 1997
- 33. Zhou K, Pedersen HK, Dawed AY, Pearson ER. Pharmacogenomics in diabetes mellitus: insights into drug action and drug discovery. Nat Rev Endocrinol 2016;12:337-346
- 34. Adler NE, Prather AA. Risk for type 2 diabetes mellitus: person, place and precision prevention. JAMA Intern Med 2015;175:1321-1322
- 35. Feng J, Glass TA, Curriero FC, Stewart WF, Schwartz BS. The built environment and obesity: a systematic review of the epidemiologic evidence. Health Place 2010:16:175-190
- 36. Mayne SL, Auchincloss AH, Michael YL. Impact of policy and built environment changes on obesity-related outcomes: a systematic review of naturally occurring experiments. Obes Rev 2015; 16:362-375
- 37. Cobb LK, Appel LJ, Franco M, Jones-Smith JC, Nur A, Anderson CA. The relationship of the local food environment with obesity: a systematic review of methods, study quality, and results. Obesity (Silver Spring) 2015;23:1331-1344
- 38. Fleischhacker SE, Evenson KR, Rodriguez DA, Ammerman AS. A systematic review of fast food access studies. Obes Rev 2011;12:e460-e471