

Association of social determinants of health with late diagnosis and survival of patients with pancreatic cancer

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Background: Pancreatic cancer disparities have been described. However, it is unknown if they contribute to a late diagnosis and survival of patients with metastatic disease. Identifying their role is important as it will open the door for interventions. We hypothesize that social determinants of health (SDH) such as income, education, race, and insurance status impact (I) stage of diagnosis of PC (Stage IV vs. other stages), and (II) overall survival (OS) in Stage IV patients.

Methods: Using the National Cancer Database, we evaluated a primary outcome of diagnosis of Stage IV PC and a secondary outcome of OS. Primary predictors included race, income, education, and insurance. Covariates included age, sex and Charlson-Deyo comorbidity score. Univariate, multivariable logistic regression models evaluated risk of a late diagnosis. Univariate, multivariable Cox proportional hazards model examined OS. 95% confidence intervals were used.

Results: 230,877 patients were included, median age of 68 years (SD 12.1). In univariate analysis, a better education, higher income, and insurance decreased the odds of Stage IV PC, while Black race increased it. In multivariable analysis, education [>93% high-school completion (HSC) *vs.* <82.4%, OR 0.96 (0.93–0.99)] and insurance [private *vs.* no, OR 0.72 (0.67–0.74)] significantly decreased the risk of a late diagnosis, whereas Black race increased the odds [*vs.* White, OR 1.09 (1.07–1.12)]. In univariate Cox analysis, having a higher income, insurance and better education improved OS, while Black race worsened it. In multivariable Cox, higher income [>\$63,333 (*vs.* <\$40,277), HR 0.87 (0.85–0.89)] and insurance [private *vs.* no, HR 0.77 (0.74–0.79)] improved OS.

Conclusions: SDH impacted the continuum of care for patients with advanced pancreatic cancer, including stage at diagnosis and overall survival.

Keywords: Survival; disparities; pancreatic cancer; social determinants of health (SDH)

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Introduction

Pancreatic adenocarcinoma (PC) is a lethal malignancy: predictions expect 60,430 new diagnoses of the disease in 2021 with 48,220 estimated deaths, and only 10% of patients are expected to live up until the 5-year mark (1). By 2030, PC is on track to be the second cause of malignancyrelated deaths in the US (2). This disease does not impact all patients equally, as healthcare outcomes for patients with PC vary by race and ethnicity as well as socioeconomic factors.

Disparities in PC have been documented by race (3-5), age (6), sex (6), income (7,8), insurance (7,8), and location (9-11). More specifically, inequities exist in how patients are cared for and treated during the early stages of the disease. For example, it is known that Black and uninsured patients have lower rates of curative intent surgery (3,4,12-14). Additionally, Black patients and racial minorities are less likely to receive adjuvant chemotherapy (13,15-16). However, it is not fully understood what impact social determinants of health (SDH), such as income, education, race, and insurance status, have on the diagnosis of PC and survival of patients with Stage IV disease, for which no curative treatments exist. Moreover, one of the limitations of some previous studies is the lack of inclusion of education as a predictive factor (8,17,18). Given that some social risk characteristics are modifiable, the study of SDH could improve the early diagnosis rate and survival of patients with pancreatic cancer.

For this reason, our study aims to evaluate if health inequities in patients with PC are more prominent before or after the diagnosis of advanced disease. We hypothesize that SDH, such as income, education, race, and insurance status, are (I) associated with stage of diagnosis of PC (Stage IV *vs.* other stages) and (II) associated with overall survival (OS) in Stage IV patients.

In our study, we sought to answer two key questions. First, what factors are associated with late-stage (Stage IV) PC diagnoses? Specifically, are a patient's race, insurance status, neighborhood income level, and/or neighborhood education level associated with late-stage diagnoses? Second, among those diagnosed with Stage IV PC, are these factors associated with survival? We present the following article in accordance with the STROBE reporting checklist (available at https://jgo.amegroups.com/article/view/10.21037/jgo-21-788/rc).

Methods

Data source and patient population

The National Cancer Database (NCDB) was queried for this retrospective observational cohort study. The NCDB encompasses 70% of hospital-based cancers diagnosed in the US. NCDB data is deidentified. Eligibility included patients 18 years and older newly diagnosed with PC in the US between January 1, 2004 to December 31, 2016. We excluded cases with more than one cancer diagnosis and cases in which the treatment was administered at a facility different from the reporting one, as this might create bias. A complete case analysis method was used for handling missing data. Cases with missing data on predictors, outcomes, and confounders were eliminated (Figure S1). 230,877 cases were selected for the final analysis. The editions 6^{th} and 7^{th} of the American Joint Commission on Cancer (AJCC) editions were used, depending on year of diagnosis. AJCC analytic stage group is a variable in NCDB. It is assigned the value of the pathologic stage. Clinical stage is used if pathologic information was not available. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013).

Outcomes

The primary outcome was late-stage diagnosis, Stage IV PC, *vs.* other stages (Stages 0-III). Staging variable was dichotomized for analysis (late or Stage IV *vs.* early or Stages 0-III). The secondary outcome was survival from time of diagnosis.

Predictor variables of interest

We used the following SDH as primary predictors: race, income, education, insurance. Race appears in the NCDB database as White, Black, and other individual twenty-seven races. These other individual race categories have fewer participants, and therefore, the race variable was defined in three categories: White, Black or other. The 2016 American Community Survey data was used to match the patient's zip code with median household income and education. Data, spanned from 2012–2016, was inflation-adjusted. Four quartiles of income were used: <\$40,227, \$40,227–\$50,353, \$50,354–\$63,332, and ≥\$63,333. Four levels of high-school graduation rate were: <82.4%, 82.5−89.1%, 89.2−93.7%, or ≥93.7%.

Control variables

Multivariable models were adjusted for age, sex, and Charlson-Deyo comorbidity score. Age was included in the analysis in ordinal categories by deciles. Sex was included as a binary variable, male *vs.* female. The Charlson-Deyo comorbidity index is calculated from diagnosis codes, applying weight for type of comorbidity, then summed for a score from 0 to 25. Values were then collapsed into four categories: a score of 0, 1, 2, or greater than 3, with higher score representing more comorbidity.

Statistical analysis

Baseline characteristics for our cohort were first summarized by early vs. late-stage (Stage 0-III vs. Stage IV). A univariate logistic regression analysis was done to analyze the impact of each variable on a diagnosis of Stage IV PC (vs. Stages 0-III). We adjusted a multivariable logistic regression model for age, sex, Charlson comorbidity index (CCI), race, insurance, income, and education. A description of the models can be found in Table S1. Finally, we did univariate and multivariable Cox proportional hazard regression models to analyze predictors associated with survival of patients with Stage IV PC. Kaplan-Meier curves were generated to illustrate differences in survival times between the categories of each of the four social determinants. Statistical analyses were done with Stata IC/SE version 16 (College Station, TX, USA). Significance was defined by P≤0.05.

Sensitivity analysis

We repeated the multivariable Cox proportional hazard regression model to document predictors associated with survival of PC patients of all stages.

Results

Patient characteristics

230,877 patients were included for the final statistical analysis. The median age was 68 years with a mean of 67.3 (standard deviation 12.1). There were 113,881 (49%) female

patients and 116,996 (51%) male patients included in the sample. Stage IV population encompassed 114,106 of the patients or a total of 49%. Only a small percentage of the included individuals lacked insurance (3%, 7,828), whereas the majority (55%, 127,094) had Medicare or private insurance (34%, 79,212). The majority of the patients were White (192,338), *vs.* Black (29,453). There was a relatively homogenous distribution of educational attainment across the four quartiles. Additional demographic characteristics can be found in *Table 1*.

Factors associated with late-stage diagnosis

In univariate analysis, education [>93% high school completion (HSC) vs. <82.4%, OR 0.93 (0.91–0.95)], income [>\$63,333 vs. <\$40,277, OR 0.94 (0.92–0.96)], and insurance [private vs. no, OR 0.70 (0.66–0.73)] significantly decreased the odds of Stage IV PC. Black race was associated with higher odds of Stage IV PC [vs. White, OR 1.11 (1.08–1.14)] (*Table 2; Figure 1*). In the multivariable analysis, education [>93% HSC vs. <82.4%, OR 0.96 (0.93–0.99)] and having insurance [private vs. no, OR 0.72 (0.67–0.74)] significantly decreased the risk of a late diagnosis, whereas Black race increased the odds of a late diagnosis [vs. White, OR 1.09 (1.07–1.12)].

Factors associated with overall survival

In univariate Cox analysis, higher income [>\$63,333 (vs. <\$40,277), HR 0.82 (0.81–0.83)], insurance [private vs. no, HR 0.77 (0.73–0.76)] and having more education (>93% HSC vs. <82.4%, HR 0.87 (0.86–0.88)] improved OS. Black race was associated with lower OS [vs. White, HR 1.03 (1.02–1.05)] (*Table 3*). In the multivariable Cox analysis, only higher income [>\$63,333 (vs. <\$40,277), HR 0.87 (0.85–0.89)] and having insurance [private vs. no, HR 0.77 (0.74–0.79)] were associated with improved OS, while Black race did not impact survival [vs. White, OR 1.00 (0.98–1.01)] (*Figures 2,3*). Older age, male sex, and a higher CCI were associated with worse survival in univariate and multivariable analysis.

In a sensitivity analysis, we evaluated the association of SDHs in the survival of patients with any stage of PC. Education, income, and insurance were associated with improved survival. Race was not significantly associated with survival (Table S2).

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Table 1 Demographic characteristics of patients with a diagnosis of pancreatic cancer

Age, deciles 18–19 53 (0%) 45 (0%) 8 (0%) 20–29 651 (0%) 435 (0%) 216 (0%) 30–39 2,623 (1%) 1,419 (1%) 1,204 (1%)	
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30–39 2,623 (1%) 1,419 (1%) 1,204 (1%)	
40–49 13,703 (6%) 6,888 (6%) 6,815 (6%)	
50–59 43,770 (19%) 21,645 (19%) 22,125 (19%)	
60–69 67,562 (29%) 34,214 (29%) 33,348 (29%)	
70–79 62,360 (27%) 32,179 (28%) 30,181 (26%)	
80–89 34,929 (15%) 17,432 (15%) 17,497 (15%)	
≥90 5,226 (2%) 2,514 (2%) 2,712 (2%)	
Sex	
Female 113,881 (49%) 59,486 (51%) 54,395 (48%)	
Male 116,996 (51%) 57,285 (49%) 59,711 (52%)	
Race	
White 192,338 (83%) 97,834 (84%) 94,504 (83%)	
African American 29,453 (13%) 14,203 (12%) 15,250 (13%)	
Other 9,086 (4%) 4,734 (4%) 4,352 (4%)	
Percent high school attainment ^a	
<82.4 48,984 (21%) 24,331 (21%) 24,653 (22%)	
82.5–89.1 60,577 (26%) 30,384 (26%) 30,193 (26%)	
89.2–93.7 65,464 (28%) 33,313 (29%) 32,151 (28%)	
>93.7 55,852 (24%) 28,743 (25%) 27,109 (24%)	
Annual household income ^b	
<\$40,227 44,704 (19%) 22,165 (19%) 22,539 (20%)	
\$40,227-\$50,353 51,215 (22%) 25,953 (22%) 25,262 (22%)	
\$50,354 - \$63,332 53,598 (23%) 27,106 (23%) 26,492 (23%)	
≥\$63,333 81,360 (35%) 41,547 (36%) 39,813 (35%)	
Insurance status	
No insurance 7,828 (3%) 3,332 (3%) 4,496 (4%)	
Private 79,212 (34%) 40,862 (35%) 38,350 (34%)	
Medicaid 14,118 (6%) 6,475 (6%) 7,643 (7%)	
Medicare 127,094 (55%) 64,634 (55%) 62,460 (55%)	
Other government 2,625 (1%) 1,468 (1%) 1,157 (1%)	

Table 1 (continued)

Table 1 (continued)

Characteristics at diagnosis	Total, N=230,877	Stage 0–III, N=116,771	Stage IV, N=114,106	
Charlson Comorbidity index $^{\circ}$				
0	150,729 (65%)	76,664 (66%)	74,065 (65%)	
1	57,205 (25%)	29,243 (25%)	27,962 (25%)	
2	14,878 (6%)	7,396 (6%)	7,482 (7%)	
≥3	8,065 (3%)	3,468 (3%)	4,597 (4%)	
		.		

^a, Percentage with high-school education from ACS 2016 matched with patient's zip code. ^b, In US dollars–Median household income from ACS 2016 matched with patient's zip-code. ^c, Charlson-Deyo Comorbidity score. ACS, American Community Survey.

Table 2 Univariate and multivariable analysis, odds of being diagnosed with Stage IV pancreatic cancer (vs. Stages 0-III)

	Univariate analysis		Multivariable analy	Multivariable analysis	
AJOO Stage IV VS. Stages 0-III	Odds ratio (95% CI)	P value	Odds ratio (95% CI)	P value	
Age (deciles)	1.00 (1.00–1.01)	0.154	1.03 (1.02–1.04)	<0.001	
Race (vs. White)					
Black	1.11 (1.08–1.14)	<0.001	1.09 (1.07–1.12)	<0.001	
Other	0.95 (0.91–0.99)	0.021	0.93 (0.90–0.98)	0.002	
Sex (vs. female)					
Male	1.14 (1.12–1.16)	<0.001	1.15 (1.13–1.17)	<0.001	
Percent high school attainment ^a					
82.5–89.1	0.98 (0.96–1.00)	0.109	1.00 (0.98–1.03)	0.981	
89.2–93.7	0.95 (0.93–0.98)	<0.001	0.98 (0.95–1.01)	0.146	
>93.7	0.93 (0.91–0.95)	<0.001	0.96 (0.93–0.99)	0.011	
Income (vs. <\$40,227) ^b					
\$40,227-\$50,353	0.96 (0.93–0.98)	0.001	0.99 (0.96–1.02)	0.45	
\$50,354-\$63,332	0.96 (0.94–0.99)	0.002	1.01 (0.98–1.04)	0.503	
>\$63,333	0.94 (0.92–0.96)	<0.001	1.01 (0.98–1.04)	0.407	
Insurance status (vs. no-insurance)					
Private	0.70 (0.66–0.73)	<0.001	0.72 (0.67–0.74)	<0.001	
Medicaid	0.87 (0.83–0.92)	<0.001	0.87 (0.83–0.92)	<0.001	
Medicare	0.72 (0.68–0.75)	<0.001	0.70 (0.66–0.73)	<0.001	
Other government	0.58 (0.53–0.64)	<0.001	0.57 (0.52–0.63)	<0.001	
Charlson Comorbidity index (vs. 0 score	∋) ^c				
1	0.99 (0.97–1.01)	0.294	0.98 (0.96–1.00)	0.054	
2	1.05 (1.01–1.08)	0.007	1.03 (1.00–1.07)	0.04	
≥3	1.27 (1.31–1.43)	<0.001	1.34 (1.28–1.40)	<0.001	

^a, percentage with high-school education from ACS 2016 matched with patient's zip code. ^b, in US dollars–Median household income from ACS 2016 matched with patient's zip-code. ^c, Charlson-Deyo Comorbidity score. AJCC, American Joint Committee on Cancer; ACS, American Community Survey.

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Figure 1 Odds of diagnosis with Stage IV pancreatic cancer (*vs.* 0-III). *Figure 1* presents in graphical form the information found in *Table 2*. OR is adjusted for age, sex, Charlson-Deyo score, insurance, race, education, and income.

Discussion

Our study showed that patients diagnosed with Stage IV PC were more likely to have lower education levels, be without insurance and be Black. Once diagnosed with metastatic disease, patients without insurance and those living in lower-income neighborhoods had shorter survival. Race and education did not have a significant impact on survival for those with Stage IV PC. These findings suggest that disparities in rates of late-stage diagnosis differ meaningfully from disparities in cancer survival. The absence of insurance was the most important factor along the continuum of PC care, from diagnosis to treatment.

Prior literature explored disparities in PC outcomes, finding that race, ethnicity, geographical location, and other socioeconomic factors impact PC survival (5,19), with unequal treatments driving much of these disparities (3,7,14,16,17). Our research builds upon this prior work by exploring the relationship between socioeconomic factors and diagnosis and survival of patients with PC with a specific focus on outcomes in the palliative setting. Furthermore, it delineates which socioeconomic factors are associated with a Stage IV diagnosis and which ones are associated with improved survival.

To our knowledge, this is the first study to show that education is an impactful factor in earlier diagnosis of PC. It is possible that more educated patients readily recognize concerning symptoms of PC such as weight loss, early satiety, and abdominal pain. These findings put an onus on public health policymakers to evaluate education tools in the community and develop strategies for disseminating knowledge about pancreatic malignancy signs and symptoms. Alternatively, higher education may be a surrogate for access to resources which facilitate followup on symptoms, risk factors or family history. While education was associated with early diagnosis, there were no differences in survival according to education for patients with metastatic disease. This suggests that despite the importance of any knowledge about the condition, once the disease is incurable, survival outcomes are superseded by other factors and resources, such as insurance and income.

Prior studies have shown that race is an independent predictor of a late diagnosis of PC and poor survival in all stages (20-24). In these studies, the survival differences by race were driven largely by differences in surgery and treatment rates in early stage or potentially curative disease settings (3,4,12,25). However, our analyses were instead focused on those with metastatic disease, eliminating the administration of curative intent treatments as mediators. Our study demonstrates that there is no impact of race on survival in the metastatic disease setting, suggesting that the racial disparities in PC survival in prior studies may be partially

Table 3 Univariate and multivariable analysis, Cox proportional hazards model for survival of patients with Stage IV pancreatic cancer

	Univariate analysis		Multivariable analy	Multivariable analysis	
Survival	Hazard ratio (95% CI)	P value	Hazard ratio (95% CI)	P value	
Age (deciles)	1.27 (1.26–1.27)	<0.001	1.25 (1.24–1.25)	<0.001	
Race (vs. White)					
Black	1.03 (1.02–1.05)	<0.001	1.00 (0.98–1.01)	0.952	
Other	0.86 (0.84–0.88)	<0.001	0.90 (0.87–0.93)	<0.001	
Sex (vs. female)					
Male	1.02 (1.01–1.02)	<0.001	1.05 (1.03–1.06)	<0.001	
Percent high school attainment ^a					
82.5–89.1	0.99 (0.98–1.00)	0.074	1.05 (1.03–1.07)	<0.001	
89.2–93.7	0.94 (0.93–0.95)	<0.001	1.04 (1.01–1.06)	<0.001	
>93.7	0.87 (0.86–0.88)	<0.001	0.99 (0.97–1.01)	0.543	
Income (vs. <\$40,227) ^{a,b}					
\$40,227-\$50,353	0.94 (0.93–0.96)	<0.001	0.96 (0.94–0.97)	<0.001	
\$50,354-\$63,332	0.90 (0.89–0.91)	<0.001	0.92 (0.90-0.94)	<0.001	
>\$63,333	0.82 (0.81–0.83)	<0.001	0.87 (0.85–0.89)	<0.001	
Insurance status (vs. no insurance)					
Private	0.74 (0.73–0.76)	<0.001	0.77 (0.74–0.79)	<0.001	
Medicaid	0.94 (0.92–0.97)	<0.001	0.93 (0.89–0.96)	<0.001	
Medicare	1.11 (1.09–1.15)	<0.001	0.84 (0.82–0.87)	<0.001	
Other government	0.85 (0.81–0.89)	<0.001	0.84 (0.79–0.90)	<0.001	
Charlson score (vs. 0 score) ^c					
1	1.12 (1.11–1.13)	<0.001	1.12 (1.16–1.14)	<0.001	
2	1.34 (1.32–1.37)	<0.001	1.32 (1.28–1.35)	<0.001	
≥3	1.69 (1.65–1.73)	<0.001	1.68 (1.63–1.73)	<0.001	

^a, percentage with high-school education from ACS 2016 matched with patient's zip code. ^b, in US dollars–Median household income from ACS 2016 matched with patient's zip-code. ^c, Charlson-Deyo Comorbidity score. ACS, American Community Survey.

due to treatment differences and access to care in early and potentially curative stages. Race was also not associated with differences in survival in a study done in an integrated healthcare system, when patients with PC had equal access to curative intent and palliative interventions (26).

Finally, the present study's finding that uninsured patients are most likely to have stage IV at diagnosis and have worse survival is compatible with results by other investigators (7,15,27). Some studies have shown that improving insurance coverage can mitigate disparities in PC treatment, at least at the state level, as this can increase

access to curative treatment (28). Our research builds upon prior research by demonstrating the importance of insurance coverage for those with late-stage PC across the care continuum, from initial diagnosis to treatment.

The current study results demonstrated that poor education, Black race, and being uninsured are predictors of a late diagnosis of PC. Once diagnosed and being treated with palliative goals, the two most important socioeconomic predictors of survival are income and insurance. Given that PC is expected to be the second cause of cancerassociated deaths by the end of this decade, these findings



Figure 2 Forest plot of hazard ratio for survival for stage IV pancreatic cancer. OR is adjusted for age, sex, Charlson-Deyo score, insurance, race, education, and income.



Figure 3 Kaplan-Meier curves survival estimates according to insurance and income in patients with Stage IV pancreatic cancer.

have important implications for public health policy. Our study suggests that education is an important factor at diagnosis. This is an argument in favor of communitylevel educational interventions. It shines a light on the importance of disseminating knowledge about symptoms of PC. For example, a person can easily dismiss abdominal pain, depending on the degree or chronicity. Similarly, new onset adult diabetes can also herald an underlying pancreatic pathology. Perhaps if we emphasized these points, patients and caregivers might be more empowered to seek attention sooner. Regardless of education and empowerment, our study demonstrated that insurance coverage is the most important factor in the diagnosis and survival of patients with PC, suggesting the importance of expanding access to care through regional or national health policies. Reducing disparities in cancer care will require the collective effort of many key stakeholders, including policymakers, public health officials, and healthcare providers.

Our study has several limitations. First, though the dataset was from accredited CoC institutions, there is the risk of misclassification bias when using the dataset from

the NCDB. Furthermore, because the NCDB is a hospitalbased database, the generalizability of the study may be limited and subject to some degree due to regionalization and by selection bias whereby patients needed to access a hospital in order to be captured by the NCDB. While this may introduce some limitations in our patient sample, it is unlikely to influence our results substantially, as the NCDB includes 70% of the US's hospital-based cancer population with representation by all states and Puerto Rico. Second, some of the variables, such as income and education, do not represent granular data. Instead, they are ecologic variables, assigned according to the patient's primary residence zip code, matched to the American Community Survey. Although this precludes the assignment of individual causation, it gives us an approximate idea of the importance of the different socioeconomic factors. Third, when analyzing race, there is some residual confounding in the analysis of patients of Black race. For example, some of the Black patients included in this study were of Hispanic descent, and others were not. Hispanics have a heterogeneous ancestry, including Indigenous, Caucasian, and African ancestry. Since other studies have evaluated this ethnic group (24,29), we chose to focus on the differences between Black vs. White patients irrespective of ethnicity, as we believe this would partially eliminate biases from heterogeneous ancestry. We acknowledge this is not the generally accepted standard evaluation of race/ethnicity and that even black ancestry has significant heterogeneity. Fourth, to handle missing variables, we used complete case analysis under the assumption that variables were missing at random. While this may not be an accurate assumption, complete case analysis is an accepted statistical technique to handle missing data.

Conclusions

The late diagnosis of PC is a death sentence for most patients with an increasing burden on the healthcare system and communities. Our study adds to the collective body of knowledge relating key SDH that impact the continuum of care for patients with advanced pancreatic cancer, including stage at diagnosis and overall survival.

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Footnote

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Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The NCDB file was provided to the authors for analytical purposes and was exempt from institutional review board review.

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Figure S1 Flow diagram for NCDB pancreas database cases selection. a, Missing EDUCATION variable 15,966 out of 276,395. No missing variable for Charlson. Missing INCOME 16,449. Missing INSURANCE 6,020. Missing AJCC clinical stage 27,505. Missing RACE 2,561.

Table S1 Univariate and multivariable logistic regression models to evaluate the impact of SDH on the odds of being diagnosed with Stage IV

pancreatic cancer (vs. 0-III)
Univariate models
$Y = \beta_{0+}\beta_1 X_{1(clinical stage)+}\beta_2 X_{2(age)}$
$Y = \beta_{0+}\beta_1 X_{1(clinical stage)+}\beta_2 X_{2(sex)}$
$Y = \beta_{0+}\beta_1 X_{1(clinical stage)+}\beta_2 X_{2(Charlsonscore)}$
$Y = \beta_{0+}\beta_1 X_{1(clinical stage)+}\beta_2 X_{2(race)}$
$Y = \beta_{0+}\beta_1 X_{1(clinical stage)+}\beta_2 X_{2(educationquart)}$
$Y = \beta_{0+}\beta_1 X_{1(clinical stage)+}\beta_2 X_{2(incomequart)}$
$Y = \beta_{0+}\beta_1 X_{1(clinical stage)+}\beta_2 X_{2(insurance)}$

Multivariable model

 $Y = \beta_0 + \beta_1 X_{1(clinical stage)} + \beta_2 X_{2(age)} + \beta_3 X_{3(sex)} + \beta_4 X_{4(Charlsonscore)} + \beta_5 X_{5(race)} + \beta_6 X_{6(educationquart)} + \beta_7 X_{7(incomequart)} + \beta_8 X_{8(insurance)} + \beta_8 X_{1(clinical stage)} + \beta_8$

Curring	Multivariable Analysis			
Survival	Hazard Ratio (95% CI)	P value		
Age (deciles)	1.30 (1.29–1.30)	<0.001		
Race (vs. White)				
Black	1.00 (0.99–1.02)	0.869		
Other	0.89 (0.87–0.91)	<0.001		
Sex (vs. Female)				
Male	1.04 (1.03–1.05)	<0.001		
Percent High School Attainment ^a (vs. <82.4)				
82.5-89.1	1.04 (1.03–1.06)	<0.001		
89.2-93.7	1.03 (1.02–1.05)	<0.001		
>93.7	0.98 (0.97–1.00)	0.057		
Income (vs. <\$40,227) ^{a,b}				
\$40,227-\$50,353	0.94 (0.93–0.96)	<0.001		
\$50,354-\$63,332	0.90 (0.89–0.91)	<0.001		
>\$63,333	0.84 (0.82–0.85)	<0.001		
Insurance Status (vs. no insurance)				
Private	0.78 (0.76–0.80)	<0.001		
Medicaid	0.95 (0.93–0.98)	<0.001		
Medicare	0.83 (0.81–0.86)	<0.001		
Other Government	0.84 (0.80–0.88)	<0.001		
Charlson Score (vs. 0 score)°				
1	1.10 (1.08–1.11)	<0.001		
2	1.28 (1.26–1.30)	<0.001		
≥3	1.59 1.55–1.63)	<0.001		
AJCC Stage Group				
1	2.82 (2.61–3.05)	<0.001		
2	4.10 (3.80–4.42)	<0.001		
3	6.63 (6.14–7.16)	<0.001		
4	11.52 (10.68–12.44)	<0.001		

Table S2 Multivariable Analysis. Cox Proportional Hazards model for survival of all patients with pancreatic cancer

AJCC, American Joint Committee on Cancer; ACS American Community Survey ^a, Percentage with high-school education from ACS 2016 matched with patient's zip code. ^b, In US dollars – Median household income from ACS 2016 matched with patient's zip-code. ^c, Charlson-Deyo Comorbidity score.