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Long-Term Survivors of Pancreatic Cancer: A California Population-Based Study

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Abstract

Objectives—Pancreatic cancer continues to carry a poor prognosis with survival rates that have had minimal improvement over the past four decades. We report a population-based, comprehensive analysis of long-term survivors of pancreatic adenocarcinoma diagnosed in the diverse population of California.

Methods—Data from the California Cancer Registry were used to evaluate long-term survival. A total of 70,442 patients diagnosed with pancreatic adenocarcinoma between 1988 and 2009 were identified. Logistic regression was used to identify factors associated with achieving five-year survival.

Results—The overall five-year survival was 2.5%, with minimal incremental improvements throughout the three decades. Age, stage, degree of differentiation, and surgical resection were associated with five-year survival. Furthermore, younger age and receiving care at a National Cancer Institute-designated cancer center were similarly correlated with five-year survival regardless of surgical intervention. Additionally, we identified stage, differentiation, and adjuvant chemotherapy as significant factors for long-term survival in surgically resected patients. In the unresectable patients, Asian/Pacific Islanders and Hispanics were significantly more likely to reach the five-year milestone than non-Hispanic Whites.

Conclusions—Although pancreatic cancer mortality remains high, our study highlights baseline characteristics, treatment, biological factors and ethnicity that are associated with long-term survival. These findings may serve as a springboard for further investigation.

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Keywords

Pancreatic Adenocarcinoma; Long-Term Survival; Surgical resection; Chemotherapy; Differentiation; Race/ethnicity

Introduction

Pancreatic cancer is the third leading cause of cancer-related deaths in the United States. More than 53,000 new cases and over 41,000 deaths occur annually.¹ Moreover, it is estimated to become the second-leading cause of cancer death in the United States by 2020.² Despite the various advances in multimodality treatment, long-term survival rates continue to range between 8% and 10%.^{1,3,4} Surgical resection remains the only potentially curative modality for patients with pancreatic cancer. However, only 15% to 20% of patients qualify for resection, because most tumors are locally advanced or metastatic at the time of diagnosis. Nevertheless, patients who undergo successful resection and adjuvant therapy have 5-year survival rates of approximately 20%, with a median survival time of 25 to 30 months.^{2,3,5,6}

Therefore, there is a pressing need to decipher the underlying elements responsible for the long-term survival of patients with pancreatic cancer including genetic, immune and molecular mechanisms, as well as clinical outcome and racial-socioeconomic disparities. To date, there are very few population-based survival studies analyzing long-term survival of patients with pancreatic adenocarcinoma.^{7–9} However, these analyses are limited and do not include data from the most recent decade, nor they do investigate racial-socioeconomic disparities. Racial and socioeconomic disparities have been increasingly highlighted recently, due to growing evidence of these inequalities in various aspects of the US health care system.^{10,11} Therefore, we report a population-based, comprehensive analysis of long-term survivors of pancreatic adenocarcinoma diagnosed from 1988 to 2009 in the diverse population of California.

MATERIALS AND METHODS

The California Cancer Registry (CCR) is a state-mandated registry that was established in 1985 and routinely abstracts demographic, tumor, and treatment data on all cases diagnosed in the state. The four regional registries comprising the CCR are part of the national SEER program and case ascertainment is estimated to be greater than 95% complete.¹² We obtained data from the CCR on all cases diagnosed with invasive pancreatic cancer between Jan 1, 1988 and December 31, 2009 (n = 70,442), excluding cases diagnosed on death certificate or autopsy (n = 2067), without valid dates of diagnosis or follow-up (n = 1384), without microscopically confirmed tumors (n = 15,133), with histologies other than adenocarcinoma (n = 11,839), and who were not identified as non-Hispanic White, non-Hispanic Black, Hispanic, or Asian/Pacific Islander (n = 213).

The CCR routinely performs follow-up on patient vital status through hospital follow-up as well as regular linkages to other national and state databases, including state and national vital statistics, Social Security Administration, credit agencies, and the Department of Motor

Vehicles. Vital status follow-up was complete through Dec. 31, 2014. We calculated survival time in months from date of diagnosis to date of last contact or date of death, if deceased. Cases with less than 60 months survival time who were alive at the time of last follow-up, but whose vital status had not been confirmed as of Dec. 31, 2014 (n = 257) were excluded as being lost to follow-up. Median follow-up for cases that were not deceased was 8.5 years (102.0 months, range 60.1–315.5 months). The final cohort consisted of 39,460 patients.

For each patient, CCR abstracts age, sex, race/ethnicity, anatomic site of tumor, histology, extent of disease at diagnosis, first course of cancer-directed therapy (chemotherapy, radiation, and surgery), reporting hospital, and census-block group of residence at time of diagnosis. Anatomic subsite and histology were defined by specific ICD-O-3 topography and morphology codes respectively. Adenocarcinomas were identified based on the following ICD-O-3 codes: 8140-8144, 8190, 8211, 8261-8263, 8290, 8310, 8440, 8500, 8503, 8560, 8570. Stage at diagnosis was defined using SEER Summary Stage (localized, regional, distant, unknown). SEER localized stage represents tumor confined to the pancreas, regional stage represents tumor with direct extension to adjacent organs or structures or spread to regional lymph nodes, and distant stage represents involvement of distant sites or lymph nodes. Because cancer registries generally do not collect individuallevel information on patient socioeconomic status (SES), we determined the patient's neighborhood SES using a previously described index^{13,14} that incorporates census measures of education, income, occupation, and cost of living. Each patient was assigned a quintile of neighborhood SES based on the neighborhood distribution of residents in California. Patients with a missing block group were randomly assigned to a block group within their county of residence.

The CCR identifies patients who have been diagnosed and/or treated at one of California's NCI-designated cancer centers. NCI cancer center designation can be viewed as a proxy for facilities with greater availability of specialized care and higher patient and procedure volumes.

We compared demographic and tumor characteristics between five-year survivors and those surviving less than five years, using chi square and *t*-tests, as appropriate. Kaplan Meier curves of overall mortality were compared using log rank test. Logistic regression analysis was used to identify factors associated with the binary outcome of achieving the 5-year survival milestone. In this exploratory analysis we modeled the effects with both univariate and multivariate regression of the following variables on the outcome: race/ethnicity (NH White, NH Black, Hispanic, Asian/Pacific Islander), sex, age (continuous), SES quintile, anatomic tumor location (head, body, tail, overlapping/NOS), SEER summary stage (localized, regional, distant, unknown), receipt of care at an NCI-designated cancer center (yes/no), surgical resection with curative intent (yes/no), receipt of radiation (yes/no), receipt of chemotherapy (yes/no), and year of diagnosis (1988–1995, prior to any chemotherapy approval, 1996–2003, post approval of gemcitabine in 1996, and 2004–2009, post the final result of ESPAC-1 trial). We repeated the analysis stratified on surgical resection. All analyses were performed using SAS 9.4 (SAS Institute, Cary, Ind).

RESULTS

We identified 70,442 patients who were diagnosed with pancreatic adenocarcinoma between 1988 and 2009 in the CCR. Of those patients, 39,460 meet our inclusion criteria, and 987 (2.5%) survived at least five years (Table 1). For all patients, the median age at diagnosis was 69 years, with a similar distribution by sex. The majority of patients were non-Hispanic White (70%), followed by Hispanics (14.1%), Asian/Pacific Islanders (8.4%) and non-Hispanic Black (7.9%). Approximately half of the cases (48.7%) did not have pathological differentiation documented; of the documented cases 23.3% were poorly differentiated or anaplastic tumors and 21.1% were moderately differentiated. As defined by SEER data, 7.2% of patients had localized disease, 34.9% had regional disease, and 50.4% of patients had distant disease, the remainder 7.5% had unknown SEER stage. Sixteen percent of all cases underwent surgical resection, 18 percent had radiation, and 41 percent received chemotherapy. Most patients (85%) did not receive care at an NCI designated cancer center. Yearly analysis of treatment modalities from 1988 to 2009 stratified by stage demonstrated interesting trends. The percentage of patients with regional disease who received surgery and systemic therapy increased since 1988, and the proportion of untreated patients decreased. Furthermore, the percentage of patients with distant disease who received chemotherapy and/or radiation treatment increased after the year 1996, while the percentage of non-treated patients decreased (Supplemental Figure 1).

Long-term survivors (defined as alive after five years) were significantly younger at diagnosis, median age of 66 years, as compared to those who died prior to five years (median age of 70 years), and a greater proportion were of an Asian/Pacific Islander or a Hispanic ethnic background. Furthermore, tumors of long-term survivors were more likely to be well or moderately differentiated, and the majority were located at the head of the pancreas. As expected, a greater proportion of long-term survivors had localized disease at diagnosis. Interestingly though, distant disease did not preclude long-term survival, as approximately eleven percent of long-term survivors presented with distant disease.

The majority of long-term survivors (75.2%) underwent surgical resection or resection combined with chemotherapy and radiation, as compared to those patients who did not survive 5 years (15%). Long-term survivors were twice as likely to have had radiation as a combined modality with chemotherapy with or without surgery (35% vs. 17%) and were more likely to have received chemotherapy (55% vs. 41%). In addition, long-term survivors were roughly twice as likely to have been seen at an NCI-designated cancer center (29% vs. 15%) and have been diagnosed in recent years. Approximately half of patients who died within five years received no cancer-directed therapy. This may reflect the era of treatment as no treatment was available prior to 1996 when gemcitabine was the first chemotherapy to be approved (Supplemental Table 1).

On further analysis using univariate logistic regression models, localized disease and surgical intervention were the most significant predictors of long-term survival (Table 2). Despite a reduction in effect size after adjustment for other covariates in the multivariate model, both stage and surgery remained the strongest predictors of long-term survival. Surgically resected patients were over 10 times more likely to reach the five-year milestone

than unresected patients (odds ratio [OR], 10.7; 95% confidence interval [CI], 8.83–13.01), although the effect of localized disease (OR, 6.41; 95% CI, 4.95–8.30) was smaller than that of surgery after the adjustment. Median survival of our cohort stratified by surgical resection was 13.7 months (95% CI, 13.1–14.0), compared to a mere 3.7 months (95% CI, 3.7–3.8) for unresectable patients (Figure 1). Furthermore, chemotherapy and radiation were also associated with five-year survival in a univariate analysis. However, after adjustment for other tumor and sociodemographic factors in a multivariate analysis, only chemotherapy maintained a significant association (OR, 1.33; 95% CI, 1.12–1.57) with long-term survival.

Well differentiated tumors were also highly associated with long-term survival (OR, 3.09; 95% CI, 2.48–3.86), however anatomic location of the tumor was not significantly associated with long-term survival after adjusting for stage. Later year of diagnosis was associated with long-term survival (OR, 1.23; 95% CI, 1.02–1.49 for diagnoses in 1996–2003 vs. 1988–1995 and OR, 1.67; 95% CI, 1.38–2.01 for patients diagnosed in 2004–2009 vs. those diagnosed in 1988–1995), as was receipt of care at an NCI-designated cancer center (OR, 1.41; 95% CI, 1.21–1.65).

Asian/Pacific Islanders and Hispanics were more likely than non-Hispanic Whites to achieve long-term survival (OR, 1.52; 95% CI, 1.23–1.88; OR, 1.24; 95% CI, 1.01–1.51, respectively). Older age was correlated with a decreased likelihood of reaching five-year survival (OR, 0.84 per 10 years; 95% CI, 0.79–0.90), but there was no significant effect of sex. Though the effect was slightly attenuated after adjustment for other factors, higher neighborhood SES was associated with a greater likelihood of being a five-year survivor (highest vs. lowest neighborhood SES, OR, 1.34; 95% CI, 1.06–1.71).

Given that surgery is one of the strongest predictors of survival, and the fact that patients with resectable disease differ clinically from those of unresectable disease, having significantly higher median survival (Fig. 1), we performed a stratified logistic regression analysis, revealing some interesting differences in the associations with long-term survival between the surgical and nonsurgical groups (Table 3). One of the most significant associations noted was for stage, which was over three times as large for localized resectable patients (OR, 12.19; 95% CI, 7.68–19.36) than localized unresectable patients (OR, 3.73; 95% CI, 2.39–5.80). Similarly, but to a lesser extent, differentiation was also associated with long-term survival. The effect of well vs. poorly-differentiated tumors was larger in resected vs unresected cases (OR, 3.29; 95% CI, 2.56–4.25 in resected patients and OR, 2.53; 95% CI, 1.62–3.95 in unresected patients). Chemotherapy was also associated with five-year survival, but only in the adjuvant setting (OR, 1.46; 95% CI, 1.19–1.80), however, radiation was not associated with long-term survival in resected patients (OR, 1.04; 95% CI, 0.85–1.28).

Similarly, later year of diagnosis (OR, 1.36; 95% CI, 1.08–1.72 for 1996–2003 vs. 1998–1995 and OR, 1.96; 95% CI, 1.55–2.48 for 2004–2009 vs. 1988–1995), neighborhood SES (OR, 1.63; 95% CI, 1.21–2.21 for highest vs. lowest quintile) and female sex (OR, 1.22; 95% CI, 1.04–1.44), were only associated with long-term survival in surgically resected cases

While race/ethnicity was not associated with five-year survival in surgically resected cases, it proved to be one of the most significant factors associated with five-year survival in unresectable disease. Asian/Pacific Islanders with unresectable disease were more than three times (OR, 3.45; 95% CI, 2.49–4.79) as likely as non-Hispanic Whites to survive over 5 years after diagnosis, and Hispanics were almost twice (OR, 1.81; 95% CI, 1.27–2.59) as likely. The magnitude of the effect of Asian race/ethnicity was on par with that of localized stage. Lastly, receiving care at an NCI-designated cancer center (OR, 1.34; 95% CI, 1.12–1.61 resected and OR, 1.63; 95% CI, 1.18–2.25 unresected) and younger age (OR per 10 years, 0.89; 95% CI, 0.83–0.96 resected and OR, 0.76; 95% CI, 0.68–0.85 unresected) were similarly associated with greater likelihood of five-year survival for both groups.

DISCUSSION

In our study, the crude overall long-term survival, defined as 5 years or longer, of patients diagnosed with pancreatic adenocarcinoma in California between 1988–2009 was 2.5%. This is similar to the percentage reported by Lambe et al¹⁵ using population-based data from a Swedish registry and Zijlstra et al from the Netherlands cancer registry,¹⁶ however, lower than the 6%-10% noted in other SEER and European databases.^{3,17} These differences may be explained by differences in methodology between the studies. While we, Lambe et al, and Zijstra et al measured long-term survival as the proportion of patients confirmed to be alive 5 years after diagnosis, Bouvier et al and Sirri et al reported 5-year relative survival, which is a theoretical measure that estimates the probability of survival at 5 years in the absence of other causes of death. Regardless of the actual percentage of five-year survival, the fact remains that in the general population, long-term survival of pancreatic adenocarcinoma has remained discouragingly low. Therefore, understanding the characteristics these individuals share may help lead to improved outcomes in the population.

Similar to previous results from the US and international population-based cancer registries, ^{1,3,18,19} we found that age, stage, degree of differentiation, and surgical resection were associated with five-year survival. Surgery is regarded as the only potentially curative option for pancreatic adenocarcinoma, and in our analysis, it was the factor most strongly associated with long-term survival. However, we also identified a subset of long-term survivors that did not undergo resection with curative intent. Because the clear majority of pancreatic cancer patients do not undergo surgery, we were particularly interested in uncovering unique associations with survivorship in this rare and understudied group of patients.

Using stratified analysis, we found that specific factors such as younger age and receiving care at an NCI-designated cancer center were similarly associated with five-year survival regardless of surgical intervention, whereas the effect of other prognostic factors differed between the two groups. In contrast to the findings in the surgically resected cohort, where clinicopathologic and treatment-related factors were more strongly associated with long-term survival, in unresected patients, race/ethnicity was almost as strongly associated with five-year survival as stage. In the unresected cohort, Asian/Pacific Islanders and Hispanics were significantly more likely to reach the five-year milestone than non-Hispanic Whites.

The improved overall survival associated with Asian race has inconsistently been reported in the literature. Using CCR data, Zell et al found that after adjusting for treatment and SES, non-Chinese Asian race was associated with a decreased hazard of death.²⁰ Gong et al reported that Asian/Pacific Islander race and receiving any active treatment at the time of diagnosis as independent factors for longer survival in SEER identified residents of San Francisco Bay Area counties.²¹ However, other SEER population-based studies have reported comparable overall survival between Asians and Whites.^{22–24} This inconsistency may relate to the lack of adjustment for SES or treatment in the latter studies. In addition, the Asian populations of California and those of other geographic areas within the US may differ in the proportion of foreign-born individuals or in the representation of the various Asian ethnic groups, which may also contribute to the inconsistent associations reported by different studies.

In contrast to Asians, previous studies have reported similar²⁰ or worse²⁵ survival outcomes for pancreatic adenocarcinoma in Hispanics compared to non-Hispanic Whites. To our knowledge, we are the first to report an association between Hispanic ethnicity and improved five-year survival outcomes. This novel finding should be confirmed by future studies.

Both the Asian and Hispanic communities in California have large immigrant populations, and it has been suggested that the survival benefit seen in these populations may be artefactual, the result of loss to follow-up due to return migration of terminally ill patients. In order to minimize this potential bias, we required confirmation of vital status at five years for all non-deceased cases. Therefore, it is unlikely that return migration could explain the improved survival we observed.

It is noteworthy that neither we nor others^{9,26} found an association with race/ethnicity after controlling for SES in surgically resected patients. While differences in long-term survival might be attributed to differential access and utilization of care among racial/ethnic groups, it appears unlikely that compared to non-Hispanic Whites, Asian and Hispanic minority patients would have improved access to care, resulting in a survival advantage. Instead, it may reflect biological differences that exist among races, which only become apparent in situations where the disease is unchecked by surgical intervention.

Dong et al previously reported differences in K-ras point mutations and p53 expression between Asian and Western populations with pancreatic carcinomas.²⁷ It is possible that our finding of a positive association of long-term survival with race, but not SES, in patients with unresectable disease reflect racial differences in biological or immunological factors or genetic-immune-environment interactions that influence the disease's aggressiveness or progression. Such differences may offer clues to the biology of the disease and warrant further investigation.

The need for a better understanding of the biology of this disease is highlighted by the finding that almost 12% of long-term survivors in our study were diagnosed with distant disease. We acknowledge that because of the limited diagnostic work-up performed in some cases, especially those that did not undergo surgical resection, some of these patients may

have been inaccurately staged, misdiagnosed or had a more favorable diagnosis such as: ampullary, peri-ampullary, or even neuroendocrine tumors. Our inability to conduct histopathologic review is one of the study's major limitations. To reduce the possibility of biasing our results by including other types of tumors or benign pancreatic masses, we required that all cases have microscopic confirmation of their pancreatic cancer diagnosis.

While stage and tumor characteristics showed the expected associations with survival in regression analysis, we recognize that the lack of histopathologic review likely resulted in some degree of misclassification, which could have potentially inflated our survival estimates. Nevertheless, it is important to note that the finding of long-term survival in the face of adverse prognostic features is not unique to our study. It is well documented that Asian/Pacific Islanders carry the lowest overall cancer incidence and death rates.¹ And it has previously been described by several other investigators in surgical case series, including those with pathologic review of specimens,^{28–30} underscoring the fact that even within the group of long-term survivors there is significant heterogeneity in tumor biology that we do not understand.

Our study has some additional limitations. The registry records limited personal and clinical information and we could not assess the effects of other potential confounders such as comorbidity, tobacco and alcohol use, family history, performance status, perioperative factors, or tumor molecular characteristics. Further, this study includes only residents of California, who may not be representative of patients in other regions.

Despite these limitations, our study has several key strengths. Ours is the largest cohort of five-year survivors of pancreatic adenocarcinoma reported to date and is the first to address long-term survival in a surgically unresected population. It is based on cancer registry data that is subject to rigorous follow-up and quality control standards. Most importantly, the population-based nature of our analysis makes our findings generalizable to a much larger population than any prior studies.

While better patient selection, surgical techniques, and postoperative treatments have benefitted patients with resectable pancreatic adenocarcinoma, it has not translated into a meaningful improvement in outcomes at the population level because too few patients present with disease amenable to resection. Until we develop strategies that lead to earlier diagnosis or therapies that are effective enough to convert the unresectable to resectable, we are unlikely to improve cure rates. The promise of targeting molecular vulnerabilities has been unfulfilled with the exception of rare patients with deficient mismatch repair (~1–2%) treated with PD-1 blockade^{31,32} and BRCA mutations (~4–5%) treated with platinum agents and experimental PARP inhibitors.^{33,34} Nevertheless, the impressive responses in these small subgroups offer proof of concept that there remain subgroups susceptible to targeted therapy and that the immune system is potent enough to be effective if there are neo-antigens to recognize. The key to these developments is achieving a better understanding of the biology and immunology of the disease, and studying this unique cohort of long-term survivors may bring us a step closer to achieving this goal.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

Acknowledgments

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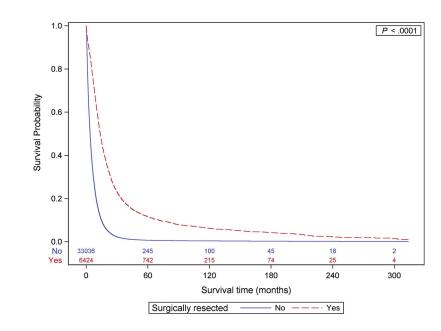


FIGURE 1. Overall Survival by Surgical Resection

TABLE 1

Characteristics of Cases With Microscopically-Confirmed Pancreatic Adenocarcinoma Diagnosed in California, 1988–2009

Kardosh et al.

| | Surviv | Survived at Least 5 Years | ars |
|--|----------------------------|------------------------------|--|
| | All n = 39,460 n (%) | N_0 n = 38,473 n (%) | $\begin{array}{c} Yes\\ n=39,460\\ n~(\%) \end{array}$ |
| Age at diagnosis | | | |
| 0-49 | 2604 (6.6) | 2493 (6.5) | 111 (11.2) |
| 50-59 | 6235 (15.8) | 6031 (15.7) | 204 (20.7) |
| 60-69 | 10,965 (27.8) | 10,673 (27.7) | 292 (29.6) |
| 62-02 | 12,974 (32.9) | 12,679 (33.0) | 295 (29.9) |
| 80+ | 2604 (6.6) | 2493 (6.5) | 111 (11.2) |
| Age at diagnosis- median years (range) | 69 (19–103) | 70 (19–103) | 66 (21–90) |
| Sex | | | |
| Male | 20,084 (50.9) | 19,603 (51.0) | 481 (48.7) |
| Female | 19,376 (49.1) | 18,870 (49.0) | 506 (51.3) |
| Race/ethnicity | | | |
| Non-Hispanic White | 27,478 (69.6) | 26,820 (69.7) | 658 (66.7) |
| Non-Hispanic Black | 3127 (7.9) | 3070 (8.0) | 57 (5.8) |
| Hispanic | 5559 (14.1) | 5408 (14.1) | 151 (15.3) |
| Asian/Pacific Islander | 3296 (8.4) | 3175 (8.3) | 121 (12.3) |
| Quintile of neighborhood SES | | | |
| 1 | 5720 (14.5) | 5602 (14.6) | 118 (12.0) |
| 7 | 7402 (18.8) | 7259 (18.9) | 143 (14.5) |
| 3 | 8371 (21.2) | 8178 (21.3) | 193 (19.6) |
| 4 | 8732 (22.1) | 8495 (22.1) | 237 (24.0) |
| S | 9235 (23.4) | 8939 (23.2) | 296 (30.0) |
| Anatomic location | | | |
| Head of pancreas | 21,574 (54.7) | 20,872 (54.3) | 702 (71.1) |
| Body of pancreas | 3822 (9.7) | 3762 (9.8) | 60~(6.1) |
| Tail of pancreas | 3902 (9.9) | 3834 (10.0) | 68 (6.9) |
| Other, NOS | 6871 (17.4) | 6755 (17.6) | 116 (11.8) |

| | Surviv | Survived at Least 5 Years | ars |
|--|----------------------------|---------------------------|----------------------------|
| | All n = 39,460 n (%) | No n = 38,473 n (%) | Yes n = 39,460 n (%) |
| Overlapping lesion | 3291 (8.3) | 3250 (8.4) | 41 (4.2) |
| Differentiation | | | |
| Well differentiated | 2714 (6.9) | 2531 (6.6) | 183 (18.5) |
| Moderately well-differentiated | 8340 (21.1) | 7939 (20.6) | 401 (40.6) |
| Poorly differentiated, undifferentiated/anaplastic | 9177 (23.3) | 8973 (23.3) | 204 (20.7) |
| Unknown | 19,229 (48.7) | 19,030 (49.5) | 199 (20.2) |
| SEER Summary stage | | | |
| Localized | 2842 (7.2) | 2592 (6.7) | 250 (25.3) |
| Regional | 13,768 (34.9) | 13,190 (34.3) | 578 (58.6) |
| Distant | 19,884 (50.4) | 19,774 (51.4) | 110 (11.1) |
| Unknown | 2966 (7.5) | 2917 (7.6) | 49 (5.0) |
| Summary treatment | | | |
| Surgery + chemo/radiation | 3312 (8.4) | 2856 (7.4) | 456 (46.2) |
| Surgery only | 3112 (7.9) | 2826 (7.3) | 286 (29.0) |
| Chemo/radiation only | 14,120 (35.8) | 13,994 (36.4) | 126 (12.8) |
| None | 18,916 (47.9) | 18,797 (48.9) | 119 (12.1) |
| Surgical resection | | | |
| No | 33,036 (83.7) | 32,791 (85.2) | 245 (24.8) |
| Yes | 6424 (16.3) | 5682 (14.8) | 742 (75.2) |
| Radiation | | | |
| No | 32,520 (82.4) | 31,880 (82.9) | 640 (64.8) |
| Yes | 6940 (17.6) | 6593 (17.1) | 347 (35.2) |
| Chemotherapy | | | |
| No | 23,226 (58.9) | 22,784 (59.2) | 442 (44.8) |
| Yes | 16,234 (41.1) | 15,689 (40.8) | 545 (55.2) |
| NCI-designated cancer center | | | |
| No | 33,552 (85.0) | 32,854 (85.4) | 698 (70.7) |
| Yes | 5908 (15.0) | 5619 (14.6) | 289 (29.3) |
| Year of diagnosis | | | |

| | Survi | Survived at Least 5 Years | ears |
|-----------|---------------------------|--|----------------------------|
| | All $n = 39,460$ n (%) | No n = 38,473 n = (0,0) | Yes n = 39,460 n (%) |
| 1988–1995 | 12,133 (30.7) | 12,133 (30.7) 11,927 (31.0) 206 (20.9) | 206 (20.9) |
| 1996–2003 | 13,626 (34.5) | 13,626 (34.5) 13,311 (34.6) 315 (31.9) | 315 (31.9) |
| 2004–2009 | 13,701 (34.7) | 13,701 (34.7) 13,235 (34.4) 466 (47.2) | 466 (47.2) |

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TABLE 2

Associations of Patient, Tumor, and Facility Characteristics With 5-Year Survival

| | Univariate Model OR (95% CI) | Multivariate Model OR (95% CI) | Multivariate Model P |
|----------------------------------|---------------------------------|-----------------------------------|----------------------|
| Age (per 10 years) | $0.76\ (0.72-0.80)$ | $0.84\ (0.79-0.90)$ | <0.0001 |
| Sex | | | 0.0641 |
| Male | Reference | Reference | |
| Female | 1.09 (0.96–1.24) | 1.14 (0.99–1.30) | |
| Race/Ethnicity | | | 0.0002 |
| Non-Hispanic White | Reference | Reference | |
| Asian/Pacific Islander | 1.55 (1.28–1.89) | 1.52 (1.23–1.88) | |
| Hispanic | 1.14(0.95 - 1.36) | 1.24 (1.01–1.51) | |
| Non-Hispanic Black | $0.76\ (0.58-0.99)$ | 0.88 (0.65–1.18) | |
| Differentiation | | | <0.0001 |
| Poorly differentiated/anaplastic | Reference | Reference | |
| Moderately differentiated | 2.22 (1.87–2.64) | 1.51 (1.26–1.80) | |
| Well-differentiated | 3.18 (2.59–3.90) | 3.09 (2.48–3.86) | |
| Unknown | 0.46 (0.38–0.56) | 1.48 (1.18–1.86) | |
| Anatomic location | | | 0.7555 |
| Head of pancreas | Reference | Reference | |
| Body of pancreas | 0.47 (0.36–0.62) | 0.89 (0.67–1.18) | |
| Overlapping, other, NOS | 0.47 (0.39–0.56) | 1.06 (0.87–1.28) | |
| Tail of pancreas | $0.53\ (0.41-0.68)$ | 1.01 (0.77–1.33) | |
| SEER Summary Stage | | | <0.0001 |
| Distant | Reference | Reference | |
| Localized | 17.34 (13.80–21.78) | 6.41 (4.95–8.30) | |
| Regional | 7.88 (6.42–9.67) | 2.40 (1.90–3.04) | |
| Unknown | 3.02 (2.15-4.24) | 4.37 (3.07–6.22) | |
| Surgery | | | <0.0001 |
| No | Reference | Reference | |
| Yes | 17.48 (15.09–20.25) | 10.72 (8.83–13.01) | |
| Radiation | | | 0.3854 |

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| | Univariate Model OR (95% CI) | Multivariate Model OR (95% CI) | Multivariate Model <i>P</i> |
|---|---------------------------------|-----------------------------------|-----------------------------|
| No | Reference | Reference | |
| Yes | 2.62 (2.29–3.00) | 1.08(0.91 - 1.28) | |
| Chemotherapy | | | 0.0010 |
| No | Reference | Reference | |
| Yes | 1.79 (1.58–2.03) | 1.33 (1.12–1.57) | |
| Quintile of neighborhood socioeconomic status | | | 0.0049 |
| 1 (lowest) | Reference | Reference | |
| 2 | 0.94 (0.73–1.20) | 0.94 (0.72–1.21) | |
| 3 | 1.12 (0.89–1.41) | 1.02 (0.79–1.30) | |
| 4 | 1.32 (1.06–1.66) | 1.19 (0.93–1.52) | |
| 5 (highest) | 1.57 (1.27–1.95) | 1.34 (1.06–1.71) | |
| Care at NCI-designated Cancer Center | | | <0.0001 |
| No | Reference | Reference | |
| Yes | 2.42 (2.10–2.78) | 1.41 (1.21–1.65) | |
| Year of diagnosis | | | <0.0001 |
| 1988–1995 | Reference | Reference | |
| 1996–2003 | 1.37(1.15–1.64) | 1.23(1.02 - 1.49) | |
| 2004–2009 | 2.04 (1.73-2.41) | 1.67 (1.38–2.01) | |

TABLE 3

Associations of Patient, Tumor, and Facility Characteristics With 5-Year Survival, Stratified by Surgical Resection

| | Unresected Total n = 33,036 | ted 13,036 | | Surgically Resected Total n = 6424 | esected 6424 | |
|---|--------------------------------|-----------------------|---------|---------------------------------------|-----------------------|---------|
| | 5-year Survivors/Non-Survivors | OR (95% CI) | P^* | 5-year Survivors/Non-Survivors | OR (95% CI) | P^* |
| Age at diagnosis per 10 years, mean (years) | 65.4/68.9 | 0.76 (0.68–0.85) | <0.0001 | 64.3/65.5 | 0.89 (0.83–0.96) | 0.0028 |
| Sex | | | | | | |
| Male | 128/16,684 | Reference | 0.8083 | 353/2919 | Reference | 0.0131 |
| Female | 117/16,107 | 0.97 (0.75–1.25) | | 389/2763 | 1.22 (1.04–1.44) | |
| Race/ethnicity | | | | | | |
| Non-Hispanic White | 126/22,790 | Reference | <0.001 | 532/4030 | Reference | 0.6424 |
| Asian/Pacific Islander | 54/2688 | 3.45 (2.49–4.79) | | 67/487 | 0.95 (0.72–1.27) | |
| Hispanic | 49/4631 | 1.81 (1.27–2.59) | | 102/777 | 1.08 (0.84–1.38) | |
| Non-Hispanic Black | 16/2682 | 1.01 (0.59–1.73) | | 41/388 | $0.84\ (0.59{-}1.19)$ | |
| Quintile of neighborhood socioeconomic status | tus | | | | | |
| 1 (lowest) | 47/4848 | Reference | 0.1652 | 71/754 | Reference | 0.0073 |
| 7 | 36/6225 | 0.62 (0.40–0.97) | | 107/1034 | 1.17 (0.84–1.62) | |
| Э | 46/7022 | 0.74 (0.49–1.13) | | 147/1156 | 1.24 (0.90–1.69) | |
| 4 | 56/7191 | 0.91 (0.60–1.36) | | 181/1304 | 1.43 (1.05–1.94) | |
| 5 (highest) | 60/7505 | $0.96\ (0.64{-}1.46)$ | | 236/1434 | 1.63 (1.21–2.21) | |
| Anatomic Site | | | | | | |
| Head of pancreas | 141/16,572 | Reference | 0.7693 | 561/4300 | Reference | 0.8552 |
| Body | 22/3484 | $0.84\ (0.53{-}1.33)$ | | 38/278 | 0.91 (0.63–1.32) | |
| Tail | 16/3403 | 0.82 (0.48–1.39) | | 52/431 | 1.11 (0.80–1.53) | |
| Overlapping, other, NOS | 66/9332 | 1.02 (0.75–1.38) | | 91/673 | 1.05 (0.82–1.34) | |
| Differentiation | | | | | | |
| Poorly differentiated/anaplastic | 44/6914 | Reference | <0.0001 | 160/2059 | Reference | <0.001 |
| Moderately differentiated | 33/5301 | 0.87 (0.55–1.37) | | 368/2638 | 1.68 (1.38–2.05) | |
| Well-differentiated | 38/1980 | 2.53 (1.62–3.95) | | 145/551 | 3.29 (2.56-4.25) | |
| Unknown | 130/18,596 | 1.00 (0.70–1.42) | | 69/434 | 2.10 (1.52-2.89) | |
| SEER summary stage | | | | | | |
| Distant | 88/19,043 | Reference | <0.001 | 22/731 | Reference | <0.0001 |

| | Unresected Total n = 33,036 | ed 3,036 | | Surgically Resected Total n = 6424 | Resected 6424 | |
|--------------------------------------|--------------------------------|------------------|--------|---------------------------------------|---------------------|---------|
| | 5-year Survivors/Non-Survivors | OR (95% CI) | P^* | 5-year Survivors/Non-Survivors | OR (95% CI) | P^* |
| Localized | 30/1906 | 3.73 (2.39–5.80) | | 220/686 | 12.19 (7.68–19.36) | |
| Regional | 88/8983 | 1.92 (1.39–2.65) | | 490/4207 | 4.14 (2.66–6.46) | |
| Unknown | 39/2859 | 3.53 (2.36–5.26) | | 10/58 | 6.95 (3.06–15.76) | |
| Radiation | | | | | | |
| No | 188/28,000 | Reference | 0.2733 | 452/3880 | Reference | 0.6811 |
| Yes | 57/4791 | 1.21 (0.86–1.71) | | 290/1802 | 1.04 (0.85–1.28) | |
| Chemotherapy | | | | | | |
| No | 130/19,753 | Reference | 0.7001 | 312/3031 | Reference | 0.0003 |
| Yes | 115/13,038 | 1.06 (0.79–1.42) | | 430/2651 | $1.46\ (1.19-1.80)$ | |
| Cancer center | | | | | | |
| No | 189/28,451 | Reference | 0.0029 | 509/4403 | Reference | 0.0013 |
| Yes | 56/4340 | 1.63 (1.18–2.25) | | 233/1279 | 1.34 (1.12–1.61) | |
| Year of diagnosis | | | | | | |
| 1988–1995 | 75/10,559 | Reference | 0.5962 | 131/1574 | Reference | <0.0001 |
| 1996–2003 | 81/11,444 | 1.07 (0.78–1.49) | | 234/2182 | 1.36 (1.08–1.72) | |
| 2004–2009 | 89/11,033 | 1.19 (0.85–1.67) | | 377/2668 | 1.96 (1.55–2.48) | |
| $^{*}_{M}$ Wald chi square P value | | | | | | |

Values in bold signify statistically significant association at P < 0.05.

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