EOPC (age 0-54) LOPC (age 55+) Total (N=713,622) (N=80,908)(N=632,714)Characteristic Ν Col % Col % Col % Ν Ν Sex Male 47,533 58.7% 317,062 50.1% 364,595 51.1% Female 33,375 41.3% 315,652 49.9% 349,027 48.9% **Race/ethnicity** Non-Hispanic White 56,007 69.2% 498,709 78.8% 77.7% 554,716 Non-Hispanic Black 13,409 16.6% 71,467 11.3% 84,876 11.9% 8,033 9.9% 40,655 6.4% 48,688 Hispanic 6.8% Non-Hispanic Asian/Pacific Islander 2,602 3.2% 17,443 2.8% 20,045 2.8% Other/Unknown 857 1.1% 4,440 0.7% 5,297 0.8% Year of diagnosis 1995-1999 11,954 14.8% 87,611 13.9% 99,565 14.0% 2000-2009 35,118 43.4% 243,150 38.4% 39.0% 278,268 2010-2018 33,836 41.8% 301,953 47.7% 335,789 47.0% Summary stage 5,242 9.5% Localized 6.5% 59,918 65,160 9.1% Regional 26,817 33.1% 202,639 32.0% 229,456 32.2% Distant 44,908 55.5% 313,350 49.5% 358,258 50.2% Unknown 3,941 4.9% 9.0% 8.5% 56,807 60,748 Grade Well differentiated; Grade I 4.1% 3,649 4.5% 25,383 4.0% 29,032 Moderately differentiated; Grade II 14,233 17.6% 91,474 14.5% 105,707 14.8% Poorly differentiated; Grade III 18.9% 15,319 96,423 15.2% 111,742 15.6% Undifferentiated; anaplastic; Grade IV 605 0.8% 3,586 0.6% 4,191 0.6% Unknown 47,102 462,950 64.9% 58.2% 415,848 65.7% Tumor site Head 39,832 49.2% 317,808 50.2% 357,640 50.1% Body 9,011 11.2% 71,334 11.3% 80,345 11.3% Tail 11,015 75,527 11.9% 86,542 13.6% 12.1% Other 21,050 168,045 189,095 26.5% 26.0% 26.6%

Supplemental Table S1. Demographic and Tumor Characteristics Stratified by Early-Onset (EOPC) and Late-Onset (LOPC) Invasive Pancreatic Cancer, 1995-2018

Supplemental Methods

Using the 1995-2018 CiNA Public Use Data Set from the North American Association of Central Cancer Registries (NAACCR), we identified incident invasive pancreatic adenocarcinoma cancer cases based on SEER Site Recode ICD-O-3/World Health Organization 2008 definitions (ICD-O-3 site codes C25.0-C25.9 and adenocarcinoma-specific histology codes) and stratified cases by EOPC (diagnosed aged 0-54) and LOPC (diagnosed age 55+). We evaluated the distribution of sex (male, female), race/ethnicity (non-Hispanic white, non-Hispanic black, Hispanic, non-Hispanic Asian/Pacific Islander, non-Hispanic American Indian, other/unknown), year of diagnosis (1995-1999, 2000-2009, 2010-2018), SEER summary stage (localized, regional, distant, unknown), grade (I, II, III, IV, unknown) and tumor site (head, body, tail, other) within EOPC and LOPC separately. Due to small sample sizes, non-Hispanic American Indian and other racial/ethnic groups were excluded from the calculation of incidence rates.

We used SEER*Stat software version 8.3.9 to obtain case distributions by demographic and tumor characteristics and calculate annual age-adjusted incidence rates (AAIR) standardized to the 2000 US Standard Population for the entire cohort and by sex and race/ethnicity. Average annual percent change (AAPC) in AAIRs was calculated using the Joinpoint Regression Program version 4.9.0.0 to analyze the time trends in AAIRs by sex and race/ethnicity. AAIRs and AAPCs were calculated for EOPC and LOPC separately.