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### An Update on Cancer in American Indians and Alaska Natives, 1999-2004

Supplement to Cancer

# Lung Cancer Incidence Among American Indians and Alaska Natives in the United States, 1999–2004

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**BACKGROUND.** Lung cancer incidence rates among American Indians and Alaska Natives (AI/ANs) in the United States have not been described well, primarily because of race misclassification and, until the 1990s, incomplete coverage of their population by cancer registries. Smoking, the predominant cause of lung cancer, is particularly prevalent among this population.

**METHODS.** Data from the National Program of Cancer Registries and the Surveillance, Epidemiology, and End Results Program were combined to estimate ageadjusted incidence rates of lung cancer during 1999 through 2004. Cases were linked to Indian Health Service (IHS) registration databases to identify AI/ANs whose race may have been misclassified. Age-adjusted rates were calculated for Contract Health Service Delivery Area (CHSDA) counties and for all counties by IHS region, and comparisons were made between AI/ANs and non-Hispanic whites (NHWs).

**RESULTS.** Among populations living in CHSDA counties, NHWs overall had higher rates of lung cancer than AI/ANs. However, the rates (per 100,000 population) among AI/ANs varied substantially between IHS regions from 14.9 (Southwest) to 87.1 (Southern Plains), 93.2 (Alaska), and 104.3 (Northern Plains). Approximately 41.6% of AI/AN lung cancer cases were diagnosed before age 65 years compared with approximately 29.8% of NHW lung cancer cases. The overall percentage stage distribution was not different between AI/ANs and NHWs. Squamous cell carcinomas were slightly more common and adenocarcinomas were less common among AI/ANs than among NHWs. Lung cancer rates were not decreasing for AI/ANs as they were for NHWs.

**CONCLUSIONS.** Data from this study clarified the need for culturally appropriate tobacco prevention and control policies and resources for AI/ANs in all regions, and especially in the Plains and Alaska. *Cancer* 2008;113(5 suppl):1168–78. *Published 2008 by the American Cancer Society.*\*

KEYWORDS: cancer, incidence, American Indian, Alaska Native, misclassification, National Program of Cancer Registries, Surveillance, Epidemiology, and End Results, United States, health disparity.

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n the United States, lung cancer is the most frequently diagnosed invasive cancer. Because it is difficult to treat, it is also the leading cause of cancer deaths, accounting for approximately 158,000 deaths each year in the United States.1 Although this cancer occurs throughout the United States population, incidence rates vary considerably between racial and ethnic populations, particularly because of the variations in tobacco use across the United States. Cigarette smoking is the most important cause of lung cancer, accounting for 90% of lung cancer deaths in males and 75%-80% of lung cancer deaths in females in the United States.<sup>2-4</sup> Exposure to other risk factors for lung cancer, such as asbestos, naturally occurring radon gas in homes, and an occupational history of uranium mining, also are contributors. 1,5-12

Several studies have demonstrated that AI/AN cancer incidence historically has been underreported with the result that the burden of cancer in this population has been described inaccurately. 13-16 This underreporting has 2 sources: First, AI/AN cancer incidence rates reported in the Surveillance, Epidemiology, and End Results (SEER) data were dominated by the large AI population ascertained by the New Mexico Tumor Registry (NMTR), which is one of the original SEER cancer registries, and often were cited as representative of all AI/AN populations in the United States. However, lung cancer incidence rates in the Southwestern AI population, as reflected in the NMTR data, are much lower than in other regions.<sup>17</sup> A second source of the underreporting is that the race of AI/ ANs often is misclassified in medical records, vital statistics, cancer registries, and other databases. Linking state cancer registry data with Indian Health Service (IHS) patient registration information can correct some underreporting because of the misclassification of race. In previous linkage studies, AI/AN mortality and incidence rates in some locations have doubled subsequent to such linkages. 18,19

The objective of this study was to reduce the misclassification of AI/AN race through linkages with analyses restricted to the IHS Contract Health Services Delivery Area (CHSDA) counties, where a larger percent of the population is AI/AN and misclassification of AI/AN race is believed to be lower than in other counties.<sup>20</sup> The cancer data used in this study are from population-based cancer registries across the United States and thereby provide an opportunity to present a more complete picture of lung cancer incidence in regional AI/AN populations.

### **MATERIALS AND METHODS**

### **Case Identification**

The cancer incidence data for this study were obtained from state and regional cancer registries throughout the United States for cases diagnosed during 1999 through 2004. All of these registries participate in either the Centers for Disease Control and Prevention's (CDC) National Program of Cancer Registries (NPCR) or the National Cancer Institute's (NCI) SEER Program,<sup>21</sup> or both. Data were included from those registries that agreed to participate and met the United States Cancer Statistics standards for high-quality data.<sup>21</sup> Only 4 states were excluded from the analysis: Kansas, Maryland, South Carolina, and Vermont. Primary cancer site and histology data were coded according to the International Classification of Diseases for Oncology (ICD-O) edition in that was use at the time of diagnosis and are converted to the third edition.<sup>22</sup> Invasive lung cancer cases were identified with the ICD-O version 3 codes C340 through C349.22 Histologies that involved hematopoietic diseases, mesothelioma, and Kaposi sarcoma were excluded (M9590-9989, 9050-9055, and 9140). For 1999 through 2000, SEER Summary Stage 1977 was used; for 2001 through 2003, SEER Summary Stage 2000 was used; and, for 2004, collaborative stage was coded, but the data were not available for analysis. For this study, stage data for 2001 through 2003 only are presented because of differences observed in comparative analyses of the 2 SEER Summary Stage systems.<sup>23,24</sup> Cell types were classified as adenocarcinoma (M8140-8389, 8401, 8408, 8410-8411, 8413, 8441, 8450, 8460, 8470, 8480-8482, 8490, 8500, 8503-8504, 8510, 8520, 8525, 8530, 8560, 8571-8574, 8576, 9070, and 9110), squamous cell carcinoma (M8050-8071, 8074-8089, and 8094), small cell carcinoma (M8041, 8044, and 8073), and large cell carcinoma (M8012-8014, 8045, and 8072).<sup>25</sup>

To obtain the most accurate race information, states worked with the IHS to link cancer registry databases with IHS patient registration data. This linkage effectively identified those individuals who may not have been classified as AI/AN in the state cancer registries but were known to be members of federally recognized tribes and enrolled in the IHS healthcare system. If an individual was not identified as AI/AN in the cancer registry but was identified as AI/AN in the IHS database, then the individual was classified as AI/AN for this analysis. For more details about the linkage methodology, see the report by Espey et al in this supplement to Cancer.<sup>26</sup> Hispanic origin was identified for all individuals in the registry databases using the North American Association of Central Cancer Registry Hispanic Identification Algo-

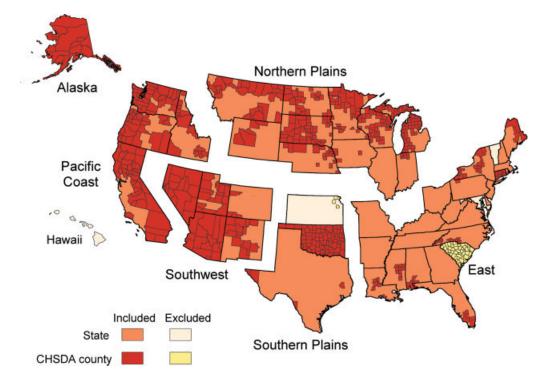


FIGURE 1. States and Contract Health Service Delivery Areas (CHSDA) counties by Indian Health Service region are illustrated.

rithm.<sup>27</sup> Individuals who were included in the AI/AN race category were included regardless of Hispanic origin. For all analyses, AI/ANs were compared with NHWs.

# Population Estimates, Regional Definitions, and Geographic Coverage

County level annual population estimates for the years 1999 through 2004 were modified in 2 ways to be used in the denominators of the rate calculations. First, since 2000, the population estimates from the US Bureau of the Census have included data from individuals who identify themselves as belonging to more than 1 racial group. The CDC National Center for Health Statistics collaborated with the Bureau of the Census to develop methods for converting the multiple race data collected into single race categories.<sup>28</sup> Second, the NCI provided additional refinements to race and county codes for use in the SEER\*Stat software program.<sup>29,30</sup> To assess regional variation in incidence, the states were grouped into 6 previously developed IHS regions (Fig. 1).26 The 'allcounties' incidence rates, for which no geographic restrictions apply, included data from registries in 46 states and the District of Columbia. For most tabulations, however, the analyses were restricted to 'Contract Health Service Delivery Area' (CHSDA) counties, which, in general, contain federally recognized tribal

lands or are adjacent to tribal lands (Fig. 1). For incidence rates restricted to CHSDA counties, data from 33 registries were included. Restricting the analysis to CHSDA counties was another method used to minimize racial misclassification. Approximately 56% of the United States AI/AN population resides in CHSDA counties. This proportion varies by IHS region (Alaska, 100%; East, 13.1%; Northern Plains, 59%; Southern Plains, 64.1%; Pacific Coast, 55.6%; Southwest, 87.5%).

### Statistical Analysis

Age-adjusted lung cancer incidence rates, expressed as the number of cases per 100,000 population and age-adjusted by 19 age groups to the 2000 United States standard population, were calculated for AI/AN and NHW populations using SEER\*Stat software version 6.3.6.<sup>30</sup> Percent distributions for stage and histology also were age-adjusted to the same standard population. For all analyses, exact counts were suppressed when the category of interest contained <6 cases.

By using the age-adjusted incidence rates, standardized rate ratios (RRs) were calculated for AI/AN populations using NHW rates for comparison. Confidence intervals (CIs) for age-adjusted rates and standardized RRs were calculated based on methods described by Tiwari et al<sup>31</sup> using SEER\*Stat 6.3.6.<sup>30</sup>

An RR >1.0 indicated that the AI/AN rate was greater than the NHW rate. Similarly, an RR <1.0 indicated an AI/AN rate less than the NHW rate. The annual percentage change (APC) was calculated for both AI/AN and NHW lung cancer incidence rates to assess trends during 1999 through 2004 (P<05). Percentage changes were calculated using 2 years for each endpoint, and the APC was not calculated if it was based on fewer than 10 cases for at least 1 year within the time interval.

### **RESULTS**

Lung cancer incidence rates in CHSDA counties varied nearly 7-fold for AI/ANs across IHS regions in the United States (Table 1). Among AI/ANs, rates were highest in the Northern Plains (104.3), Alaska (93.2), and the Southern Plains (87.1) and lowest in the Southwest (14.9). This same pattern was observed in the analysis of all counties combined, although the rates were lower and the variation was nearly 5-fold. Lung cancer incidence rates for NHWs did not exhibit the same magnitude of variation or the same pattern of regional variation; in the CHSDA counties, NHW rates were highest in the Southern Plains (82.2) and the East (74.2). All subsequent presentations of data refer to CHSDA county data unless noted otherwise.

Among the populations living in CHSDA counties, AI/ANs had significantly higher rates (P < .05) of lung cancer than NHWs for both males and females in the Northern Plains and Alaska and for females in the Southern Plains (Table 1). In the Southern Plains, AI/AN males had lung cancer incidence rates that were approximately the same as those for NHWs, although that rate was significantly higher than the national NHW rate. In contrast, incidence rates for AI/ANs in the Pacific Coast, East, and Southwest regions all were significantly lower than the rates for NHWs in the same regions for both males and females (Table 1, Fig. 2). 32 From 1999 through 2004, male NHW lung cancer incidence rates decreased significantly for all regions combined, whereas male AI/AN rates remained approximately level (Table 2). Over the same period, rates among AI/AN females and NHW females did not change.

The highest rates among AI/ANs for each age group were in the Northern Plains, Alaska, and the Southern Plains (Table 3). Similar regional patterns were not observed for NHWs, and their highest rates were observed among those living in the Southern Plains and the East. Like for many other forms of cancer, rates increased with age, and the highest rates occurred in the oldest age group (aged

 $\geq$ 65 years). Among individuals aged  $\geq$ 50 years, lung cancer incidence rates were significantly higher among AI/ANs than among NHWs in the Northern Plains and Alaska. Also, among individuals aged  $\geq$ 50 years, the rates were significantly lower among AI/ANs than among NHWs in the Southwest, East, and Pacific Coast regions. AI/ANs were diagnosed with lung cancer at younger ages than NHWs. Among AI/ANs, 58.5% of cases were aged  $\geq$ 65 years at diagnosis compared with 70.2% of cases among NHWs.

Table 4 displays an analysis of the lung cancer incidence by disease stage at diagnosis among AI/ ANs and NHWs for CHSDA counties in the 6 IHS regions. For both populations across all regions, approximately 44% of cases were diagnosed at a distant stage. In the Southwest, AI/ANs had the highest percentage of distant-stage disease (53.3%) and the lowest percentage of local-stage disease (11.3%). Although AI/ANs in the Southern Plains had among the higher lung cancer incidence rates, AI/ANs in the same region had the lowest percentage of cases diagnosed with distant-stage disease (40.5%). For all regions combined, AI/AN females were more likely than AI/AN males to have their lung cancer diagnosed at a local stage and to have less distant-stage disease (data not shown). Among NHWs, the highest proportion of distant-stage disease was observed in the Pacific Coast and Alaska regions; the percentage stage distributions did not differ appreciably by sex (data not shown).

A comparison of the percentage distribution of histologic type among microscopically confirmed lung and bronchus cancer in cases between AI/ANs and NHWs in CHSDA counties is presented in Table 5. For all regions combined, AI/ANs had relatively less adenocarcinoma (29.2%) and slightly more squamous cell histology (24.0%) than NHWs (37.1% and 20.4%, respectively), except in the Southern Plains and Southwest, where the histology distributions for the 2 populations were similar.

### **DISCUSSION**

This is the most comprehensive study of the incidence of lung cancer among the AI/AN populations in the United States to date. Combining NPCR and SEER data for broad population coverage and using linkages to improve on the misclassification of race among AI/ANs provided the opportunity to describe both regional and national estimates of lung cancer incidence for AI/ANs. Our findings confirmed the dramatic regional differences in lung cancer rates that have been reported in studies using death certificates. <sup>16</sup> Lung cancer incidence rates in AI/ANs var-

Invasive Lung and Bronchus Cancer Incidence Rates by Indian Health Service Region and Sex for American Indians/Alaska Natives and Non-Hispanic Whites: United States, 1999-2004<sup>a</sup>

				CHSDA Counties	ş					All C	All Counties		
IHS Region	Sex	AI/AN Count	AI/AN Rate <sup>b</sup>	95% CI for AI/AN Rate	NHW Rate <sup>b</sup>	RR (AI/AN:NHW)°	95% CI for RR	AI/AN Count	AI/AN Rate <sup>b</sup>	95% CI for AI/AN Rate	NHW Rate <sup>b</sup>	RR (AI/AN:NHW)°	95% CI for RR
Northern Plains	Both Sexes	719	104.3	96.4-112.7	65.5	1.59 <sup>d</sup>	1.47-1.72	951	81.9	76.4-87.7	9.69	$1.18^{d}$	1.10-1.26
	Males	349	119.8	106.3-134.3	84.6	$1.42^{d}$	1.25-1.59	467	95.9	86.4-106.1	89.7	1.07	0.96-1.18
	Females	370	93.8	84.1-104.2	51.1	1.83 <sup>d</sup>	1.64-2.04	484	72.6	2.62-0.99	55.3	1.31 <sup>d</sup>	1.19-1.44
Alaska <sup>e</sup>	Both sexes	326	93.2	82.9-104.2	71.6	$1.30^{d}$	1.14-1.48	326	93.2	82.9-104.2	71.6	$1.30^{d}$	1.14-1.48
	Males	185	115.3	97.9-134.7	85.0	$1.36^{d}$	1.13-1.62	185	115.3	97.9-134.7	85.0	$1.36^{d}$	1.13-1.62
	Females	141	75.4	63.1-89.1	2.09	$1.24^{d}$	1.02-1.50	141	75.4	63.1-89.1	2.09	$1.24^{d}$	1.02-1.50
Southern Plains	Both sexes	1131	87.1	82.0-92.4	82.2	1.06	1.00-1.13	1227	69.3	65.4-73.5	77.3	$^{0.90^{d}}$	0.85-0.95
	Males	620	111.0	101.9-120.5	108.5	1.02	0.94 - 1.11	999	82.8	79.0-93.1	100.7	$0.85^{d}$	0.78-0.92
	Females	511	6.69	63.9-76.3	9.79	$1.12^{d}$	1.02-1.22	295	57.2	52.5-62.2	0.09	0.95	0.87-1.04
Pacific Coast	Both sexes	902	51.8	47.5-56.4	70.1	$0.74^{d}$	0.68-0.80	755	35.8	33.1-38.6	8.99	$0.54^{ m d}$	0.50-0.58
	Males	293	57.6	50.6-65.3	82.5	$0.70^{d}$	0.61-0.79	373	40.2	35.9-44.9	78.3	$0.51^{d}$	0.46 - 0.57
	Females	312	48.0	42.6-53.9	6.09	0.79 <sup>d</sup>	0.70-0.88	382	32.7	29.4-36.3	58.4	$0.56^{\mathrm{d}}$	0.50-0.62
East	Both sexes	144	46.8	39.2-55.4	74.2	$0.63^{d}$	0.53-0.75	639	28.4	26.1-30.8	75.8	$0.37^{d}$	0.34-0.41
	Males	71	51.0	39.0-65.3	91.5	$0.56^{\mathrm{d}}$	0.43-0.71	379	37.8	33.8-42.2	8.96	$0.39^{d}$	0.35-0.44
	Females	73	43.5	33.9-54.7	61.7	$0.71^{d}$	0.55-0.89	260	21.0	18.5-23.8	60.4	$0.35^{d}$	0.31-0.39
Southwest	Both sexes	240	14.9	13.0-17.0	9.59	0.23 <sup>d</sup>	0.20-0.26	268	15.0	13.2-16.9	59.2	$0.25^{d}$	0.22-0.29
	Males	144	21.2	17.7-25.1	77.3	$0.27^{\rm d}$	0.23-0.33	161	21.1	17.8-24.8	6.07	$0.30^{d}$	0.25-0.35
	Females	96	10.4	8.4-12.7	56.3	$0.18^{d}$	0.15-0.23	107	10.5	8.6-12.7	50.1	$0.21^{d}$	0.17-0.25
Total	Both sexes	3165	57.4	55.4-59.5	70.3	$0.82^{\rm d}$	0.79-0.85	4166	43.8	42.4-45.2	72.4	$0.61^{d}$	0.59 - 0.62
	Males	1662	69.5	66.0-73.2	85.9	$0.81^{ m d}$	0.77-0.85	230	53.7	51.3-56.2	91.4	0.59 <sup>d</sup>	0.56 - 0.61
	Females	1503	48.5	46.0-51.1	9.85	0.83 <sup>d</sup>	0.78-0.87	1936	36.5	34.8-38.2	58.5	0.62 <sup>d</sup>	0.60-0.65

Source: Cancer registries in the Centers for Disease Control and Prevention's National Program of Cancer Registries (NPCR) and or the National Cancer Institute's Surveillance, Epidemiology, and End Results (SEER) Program. CHSDA indicates Contract Health Service Delivery Area; IHS, Indian Health Service, AI/AN, American Indians/Alaska Natives, NHW, non-Hispanic whites; CI, confidence interval.

Percentage regional coverage of AI/AN in CHSDA counties compared with AI/AN in all counties: Alaska, 100%; East, 13.1%; Northern Plains, 59.0%; Southern Plains, 64.1%; Pacific Coast, 55.6%; Southwest, 67.5%.

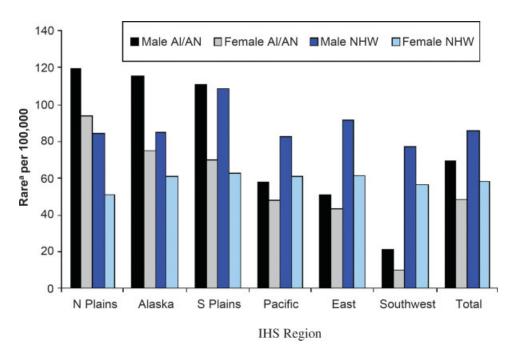
AI/AN race is reported by NPCR and SEBR registries or through linkage with the IHS patient registration database. AI/AN persons of Hispanic origin are included.

b Rates are per 100,000 persons and are age-adjusted to the 2000 U.S. standard population (19 age groups, Census P25-1130; see Day 1996<sup>32</sup>).

c RRs were calculated in SEER\*Stat prior to rounding of rates and may not equal the RRs calculated from rates presented in the table.

<sup>&</sup>lt;sup>d</sup> The RR is statistically significant (P < .05).

Years of data and registries used. 1999-2004 (41 states and the District of Columbia): Alakaka, \*Alabama, \*Arkansas, Arizona, \*Colorado, \*Connecticut, \* the District of Columbia, Delaware, Florida, \*Georgia, Hawaii, Jowa, \*Idaho, \*Illinois, Indiana, \*Kahama, \*Arkansas, Arizona, \*Colorado, \*Connecticut, \* the District of Columbia, Delaware, Florida, \*Georgia, Hawaii, Jowa, \*Idaho, \*Illinois, Indiana, \*Kahama, \*Arkansas, Arizona, \*Colorado, \*Connecticut, \* the District of Columbia, Delaware, Florida, \*Georgia, Hawaii, Jowa, \*Idaho, \*Illinois, Indiana, \*Kahama, \*Illinois, Indiana, \*Illinois, Illinois, Massachusetts, \*Maine, \*Michigan, \* Missouri, Montana, \*North Carolina, \*North Carolina, \*North Carolina, \*North Hampshire, New Hampshire, New Mexico, \*Nevada, \*New York, \*Ohio, Oklahoma, \*Oregon, \*Pennsybaia, \*Rhode Island, \*Texas, \*Ulah, \*Washington, \*Wisconsin, \*West Virginia and Wyoning\*; 1999 and 2002-2004; North Dakota\*; 2001-2004; South Dakota\*; 2003-2004; Mississippi\* and Virginia; 2004; Tennessee (asterisks indicate states with at least 1 county designated as a CHSDA). e Rates and RRs for Alaska in the CHSDA Counties section are the same as those in the All Counties section, because all counties in Alaska are CHSDA counties.



**FIGURE 2.** Lung and bronchus cancer incidence rates for American Indians and Alaska Natives (Al/AN) and non-Hispanic whites (NHW) by Indian Health Service (IHS) region and Sex in Contract Health Service Delivery Area (CHSDA) counties in the United States from 1999 to 2004. <sup>a</sup>Rates are age-adjusted to 2000 United States standard population (19 age groups; Census P25-1130; see Day, 1996<sup>32</sup>). N Plains indicates Northern Plains; S Plains, Southern Plains. Source: cancer registries in the Centers for Disease Control and Prevention's National Program of Cancer Registries and/or the National Cancer Institute's Surveillance, Epidemiology, and End Results Program.

ied 7-fold across IHS regions (in CHSDA counties), with the highest rates observed in the Northern and Southern Plains and Alaska and the lowest rates observed in the Southwest. AI/AN males and females were diagnosed with lung cancer at a younger age than NHWs, but the distribution of disease stage did not differ between the 2 populations. Adenocarcinoma was less common and squamous cell histology was slightly more common among AI/ANs than among NHWs.

Cigarette smoking and exposure to environmental tobacco have been linked directly to lung cancer and many other diseases, including heart disease, stroke, and multiple cancers, 2,3,33 that translate into multibillion dollar medical and lost productivity costs in the United States. 4 AI/ANs have the highest prevalence of daily cigarette use of any population in the United States. The prevalence of tobacco use varies greatly among AI/ANs by region. Thus, regional lung cancer rates also can be expected to vary; and, similar to the current study, such variations were observed in earlier incidence and mortality studies. 13-16,37

Tobacco use survey information must be interpreted with caution, because many American Indians use tobacco for traditional or ceremonial purposes

TABLE 2 Invasive Lung and Bronchus Cancer Incidence Annual Percentage Change by Year of Diagnosis and Indian Health Service Region for American Indians/Alaska Natives and Non-Hispanic Whites in Contract Health Service Delivery Area Counties: United States, 1999-2004

	Both	Sexes	Ma	les	Fem	ales
IHS Region	AI/AN APC	NHW APC	AI/AN APC	NHW APC	AI/AN APC	NHW APC
Northern Plains	-4.5	-0.5	-3.1	-0.9	-5.9	-0.1
Alaska	5.6	-2.6	2.0	-0.7	10.8	-5.2
Southern Plains	1.6	-0.7	2.1	-2.2	0.2	1.1
Pacific Coast	0.9	$-1.8^{a}$	2.6	$-2.8^{a}$	0.0	$-0.8^{a}$
East	-10.8	$-0.8^{a}$	$-16.0^{a}$	$-1.8^{a}$	-8.2	0.2
Southwest	-5.1	-1.1	-2.8	$-2.1^{a}$	-7.5	-0.1
Total	-0.2	-1.2 <sup>a</sup>	0.1	$-2.1^{a}$	-0.6	-0.3

Source: Cancer registries in the Centers for Disease Control and Prevention's National Program of Cancer Registries and/or the National Cancer Institute's Surveillance, Epidemiology, and End Results Program.

IHS indicates Indian Health Service; AI/AN, American Indians/Alaska Natives; APC, annual percentage change; NHW, non-Hispanic whites.

and may or may not also be habitual smokers. 38,39 Tobacco has been a key element in the ceremonial practices of many American Indian people from pre-

<sup>&</sup>lt;sup>a</sup> The APC is significantly different from zero (P < .05)

TABLE 3
Invasive Lung and Bronchus Cancer Incidence Rates by Age and Indian Health Service Region for American Indians and Alaska Natives and Non-Hispanic Whites in Contract Health Service Delivery Area Counties: United States, 1999-2004

		<50 Ye	ars			50-64	Years			≥65 Y	ears	
IHS Region	Count	% of Cases <sup>a</sup>	Rate <sup>b</sup>	95% CI	Count	% of Cases <sup>a</sup>	Rate <sup>b</sup>	95% CI	Count	% of Cases <sup>a</sup>	Rate <sup>b</sup>	95% CI
AI/AN												
Northern Plains	48	6.7	5.6	4.1-7.4	261	36.3	179.3 <sup>c</sup>	158.2-202.4	410	57.0	580.8 <sup>c</sup>	524.2-641.9
Alaska	19	5.8	4.6	2.8-7.1	113	34.7	163.2 <sup>c</sup>	134.4-196.2	194	59.5	517.0°	445.0-597.5
Southern Plains	107	9.5	8.9	7.3-10.7	376	33.2	157.4	141.9-174.1	648	57.3	451.3	416.9-487.8
Pacific Coast	30	5.0	$2.0^{d}$	1.3-2.8	221	36.5	$83.6^{\mathrm{d}}$	72.9-95.4	354	58.5	299.4 <sup>d</sup>	268.2-333.2
East	13	9.0	3.6	1.9-6.1	46	31.9	72.8 <sup>d</sup>	53.2-97.0	85	59.0	263.3 <sup>d</sup>	209.6-326.7
Southwest	13	5.4	$0.6^{\mathrm{d}}$	0.3-1.1	68	28.3	$20.9^{d}$	16.2-26.5	159	66.3	$89.7^{\mathrm{d}}$	76.1-105.0
Total	230	7.3	$3.7^{\mathrm{d}}$	3.2-4.2	1085	34.3	$98.0^{\mathrm{d}}$	92.3-104.0	1850	58.5	317.0 <sup>d</sup>	302.4-332.0
NHW												
Northern Plains	1686	5.0	4.9	4.7-5.2	8809	26.2	114.2	111.8-116.6	23,131	68.8	354.8	350.3-359.4
Alaska	111	8.8	4.6	3.8-5.5	450	35.9	108.5	98.6-119.1	694	55.3	411.9	381.2-444.4
Southern Plains	808	4.8	6.7	6.2-7.2	4521	27.1	148.8	144.5-153.3	11,333	68.0	435.3	427.3-443.4
Pacific Coast	2720	4.1	4.3	4.1-4.5	16,144	24.4	112.0	110.2-113.7	47,311	71.5	397.3	393.7-400.9
East	2358	4.7	5.9	5.6-6.1	12,572	25.0	130.8	128.5-133.1	35,331	70.3	398.1	393.9-402.3
Southwest	1095	3.8	4.3	4.1-4.6	7451	25.6	111.9	109.4-114.5	20,549	70.6	361.8	356.8-366.8
Total	8778	4.5	4.9	4.8-5.0	49,947	25.3	119.3	118.2-120.3	138,349	70.2	386.7	384.7-388.8

Source: Cancer registries in the Centers for Disease Control and Prevention's National Program of Cancer Registries and/or the National Cancer Institute's Surveillance, Epidemiology, and End Results Program. IHS indicates Indian Health Service; CI, confidence interval; AI/AN, American Indians/Alaska Natives; NHW, non-Hispanic whites.

TABLE 4
Invasive Lung and Bronchus Cancer Incidence, Percentage Distribution by Summary Stage and Indian Health Service Region for American Indians and Alaska Natives and Non-Hispanic Whites in Contract Health Service Delivery Area Counties: United States, 1999-2003<sup>a</sup>

		Lo	calized			Re	gional			D	istant			Un	staged	
IHS Region	Count	Rate <sup>b</sup>	95% CI	% of Cases <sup>c</sup>	Count	Rate <sup>b</sup>	95% CI	% of Cases <sup>c</sup>	Count	Rate <sup>b</sup>	95% CI	% of Cases <sup>c</sup>	Count	Rate <sup>b</sup>	95% CI	% of Cases <sup>c</sup>
AI/AN																
Northern Plains	64	17.4	13.2-22.4	16.8	109	26.3	21.4-32.0	25.4	170	44.4	37.6-51.9	42.9	45	15.3	11.0-20.5	14.8
Alaska	24	13.3	8.3-19.9	14.4	43	24.0	17.2-32.5	25.9	79	41.8	32.7-52.4	45.1	19	13.6	8.0-21.0	14.7
Southern Plains	98	14.9	12.1-18.2	17.0	135	20.0	16.7-23.7	22.9	239	35.4	30.9-40.2	40.5	106	17.1	14.0-20.8	19.6
Pacific Coast	50	8.1	5.9-10.8	14.4	72	12.1	9.3-15.4	21.5	169	28.1	23.7-32.9	49.9	41	8.1	5.7-11.0	14.4
East	16	10.0	5.6-16.2	19.4	19	11.4	6.7-18.0	22.1	33	21.4	14.5-30.2	41.5	11	8.7	4.3-15.3	16.9
Southwest	15	1.7	0.9-2.9	11.3	21	2.4	1.4-3.6	16.0	65	8.0	6.1-10.2	53.3	25	3.0	1.9-4.4	20.0
Total	267	9.4	8.3-10.6	16.0	399	13.5	12.2-15.0	22.9	755	26.2	24.2-28.2	44.5	247	9.8	8.6-11.2	16.6
NHW																
Northern Plains	3156	12.1	11.7-12.6	18.4	4345	16.7	16.2-17.2	25.4	6864	26.4	25.8-27.0	40.2	2786	10.4	10.0-10.8	15.8
Alaska	95	11.2	8.9-13.8	15.6	141	15.7	13.0-18.7	21.8	318	34.1	30.1-38.4	47.4	83	11.1	8.7-13.8	15.4
Southern Plains	1535	15.1	14.3-15.8	18.1	1995	19.7	18.8-20.5	23.6	3513	34.7	33.6-35.9	41.6	1438	14.1	13.4-14.8	16.9
Pacific Coast	5386	11.4	11.1-11.7	16.4	7679	16.3	15.9-16.6	23.4	16,529	34.9	34.3-35.4	50.1	3414	7.0	6.8-7.3	10.1
East	4315	12.7	12.4-13.1	17.2	5655	16.9	16.4-17.3	22.8	10,823	32.1	31.5-32.7	43.4	4331	12.3	11.9-12.7	16.6
Southwest	2219	9.9	9.5-10.3	14.9	2747	12.3	11.8-12.7	18.6	5,900	26.5	25.8-27.2	40.0	3956	17.7	17.1-18.2	26.7
Total	16,706	11.9	11.7-12.1	16.9	22,562	16.1	15.9-16.3	22.9	43,947	31.2	31.0-31.5	44.3	16,008	11.1	11.0-11.3	15.8

Source: Cancer registries in the Centers for Disease Control and Prevention's National Program of Cancer Registries and/or the National Cancer Institute's Surveillance, Epidemiology, and End Results (SEER) Program.

<sup>&</sup>lt;sup>a</sup> Percentages may not add to 100% because of rounding.

b Rates are per 100,000 persons and are age-adjusted to the 2000 U.S. standard population (19 age groups; Census P25-1130; see Day 1996<sup>32</sup>).

 $<sup>^{\</sup>rm c}$  The AI/AN rate is statistically significantly higher than the NHW rate (P<05).

 $<sup>^{\</sup>mathbf{d}}$  The AI/AN rate is statistically significantly lower than the NHW rate (P<05).

IHS indicates Indian Health Service; CI, confidence interval; AI/AN, American Indians/Alaska Natives; NHW, non-Hispanic whites.

<sup>&</sup>lt;sup>a</sup> SEER Summary Stage 2000 was used.

b Rates are per 100,000 persons and are age-adjusted to the 2000 U.S. standard population (19 age groups; Census P25-1130; see Day 1996<sup>32</sup>).

c Percentages are age-adjusted to the 2000 U.S. standard population; percentages may not add to 100% because of rounding.

Percentage Distribution of Histologic Type of Invasive Lung and Bronchus Cancer Among Microscopically Confirmed Cases for American Indians/Alaska Natives and Non-Hispanic Whites by Indian Health Service Region in Contract Health Service Delivery Counties: United States, 1999-2004

		Adenocarcinoma <sup>a</sup>	cinoma <sup>a</sup>		nbs	Squamous Cell	U Carcinon	1a <sup>b</sup>	S	Small Cell Carcinoma	)arcinoma <sup>c</sup>		Ľ	Large Cell Ca	Carcinoma <sup>d</sup>		Othe	Other Malignan	t Histolog	ese
	AI,	(AN	NHW	IW	AI/,	AN	NHW	M.	AI/,	AN	NHW	M	AI/£	N	NHW	M	AI/AN	N	NHW	W
IHS Region	Count	% of Cases <sup>f</sup>	Count	% of Cases <sup>f</sup>	Count	% of Cases <sup>f</sup>	Count	% of Cases <sup>f</sup>	Count	% of Cases <sup>f</sup>	Count	% of Cases <sup>f</sup>	Count	% of Cases <sup>f</sup>	Count	% of Cases <sup>f</sup>	Count	% of Cases <sup>f</sup>	Count	% of Cases <sup>f</sup>
Northern Plains	182	27.0	10,529	36.0	162	26.4	6537	22.3	120	18.8	4616	15.9	36	6.2	1634	5.6	134	21.6	5943	20.2
Alaska	77	25.1	453	39.8	9/	29.0	232	22.4	48	16.4	172	15.0	11	3.8	20	4.2	72	25.7	207	18.5
Southern Plains	306	31.4	4339	31.3	199	21.4	3119	22.3	140	14.1	2154	15.6	85	9.1	1272	9.5	224	24.1	3021	21.7
Pacific Coast	148	28.2	21,933	38.4	112	22.8	11,253	19.7	93	17.7	2987	13.8	28	5.4	2735	4.8	136	25.9	13,398	23.3
East	36	28.1	17,240	39.3	35	27.8	9888	20.0	19	15.7	5938	13.6	6	8.5	2266	5.2	25	20.1	9722	22.0
Southwest	89	34.2	8778	35.0	41	22.5	4947	9.61	32	14.2	3512	14.0	6	5.0	1372	5.5	47	24.2	6527	26.0
Total	817	29.2	63,272	37.1	625	24.0	34,974	20.4	452	16.1	24,259	14.2	178	6.9	9329	5.6	638	23.8	38,818	22.7

Source: Cancer registries in the Centers for Disease Control and Prevention's National Program of Cancer Registries and/or the National Cancer Institute's Surveillance, Epidemiology, and End Results Program (for the states included, see Table 1). AI/AN indicates American Indians/Alaska Natives; NHW, non-Hispanic whites; IHS, Indian Health Service.

and brindings histology codes 8140-8389, 8401, 8408, 8410-8411, 8413, 8441, 8450, 8460, 8470, 8480-8482, 8490, 8503-8504, 8510, 8520, 8525, 8530, 8560, 8571-8574, 8576, 9070, and 9110.

 $^{\rm b}$  Includes histology codes 8050-8071, 8074-8089, and 8094.

<sup>c</sup> Includes histology codes 8041, 8044, and 8073.

<sup>d</sup> Includes histology codes 8012-8014, 8045, and 8072.

e Included histology codes 8000-8011, 8015-8040, 8042-8043, 8046-8049, 8096-8093, 8096-8093, 8095-8139, 8350-8400, 8412, 8414-8440, 8442-8449, 8461-8469, 8471-8479, 8483-8489, 8491-8499, 8501-8502, 8505-8509, 8511-8519, 8521-8524, 8526-8529, 8531-8559, 8561-8570, 8575, 8577-9069, 9071-9109, and 9111-9989.

f Percentages are age-adjusted to the 2000 U.S. standard population; percentages may not add to 100.0% because of rounding.

Columbian times. Many Native stories emphasize the sacredness of the plant and its powers to heal if used properly and to harm if used improperly. Traditional tobacco mixes may include the Nicotiana plant, kinnick-kinnick, and other herbs. However, commercial tobacco products, by virtue of convenience, are used sometimes for ceremonial functions. Inhalation of tobacco smoke during ceremonial use typically is brief and does not occur every day; therefore, it is unlikely to lead to lung cancer.

Data from 15 years of Behavioral Risk Factor Surveillance System (BRFSS) surveys indicate that AI/AN current smoking rates have not been declining. 36,38-40 According to aggregate data from the 2000 through 2006 BRFSS, AI/AN respondents were more likely to be currently smoking cigarettes (31.1%) than the NHW population (22.8%). Current cigarette smoking was highest in the Northern Plains (40.2%) and Alaska (40%) and was lowest in the Southwest (21.1%). For all racial and ethnic groups, data on current cigarette smoking from the 2006 National Health Interview Survey show the highest rate among AI/ANs (26.9%), followed by NHWs (22.6%), Native Hawaiian/Pacific Islanders (22.5%), African Americans (21.8%), Hispanics (14.5%), and Asians (10.7%).

Although it is estimated that tobacco is a causal factor in nearly 90% of lung cancers, other occupational and environmental exposures also are known lung carcinogens. These include asbestos, radon, arsenic, and uranium mining.1 According to a 1999 Institute of Medicine report, an estimated 21,800 new diagnoses of lung cancer are related to radon (14%) each year in the United States, which makes radon the second leading cause of lung cancer deaths after cigarette smoking.41 Other studies have suggested a synergistic relation between radon exposure and cigarette smoking that may increase the risk of lung cancer to 6 to 10 times higher than among nonsmokers. 11,12,42 Radon gas occurs naturally in small amounts in soils and rocks throughout the United States. Radon can become trapped in homes and other structures, creating the potential for elevated exposure.<sup>43</sup> The Environmental Protection Agency (EPA) has established 4 pico Curies per liter (pCi/L) as the threshold above which action should be taken to alter a structure to reduce radon exposure.44 EPA maps of radon zones indicate that homes in the part of the country with a high proportion of AI/ANs contain high (4 pCi/L) or moderate (2 pCi/L) levels of radon, which may contribute to some cases of lung cancer among AI/ANs.45 Another form of exposure to radon is in the process of uranium mining. This occupational exposure has long been documented as a strong causal factor in the development of lung cancer. In the Southwest, many AI/AN males worked in uranium mines in the 1950s and 1960s; it has been demonstrated that this is an important cause of lung cancer in a population that historically has had a very low prevalence of smoking.<sup>5,6</sup> The Southwest region has had the lowest lung cancer incidence rates, despite the extensive exposure to uranium mining that occurred in that region.

Other known risk factors for lung cancer may include tuberculosis and chronic obstructive pulmonary disease. Exposure to arsenic, chromium, nickel, aromatic hydrocarbons, and ethers, most commonly in an occupational setting, also may increase the risk of lung cancer. 47

The IHS patient registration database contains records of individuals who are members of federally recognized tribes, and linking these data with SEER and NPCR registries corrected some of the misclassification of race for AI/ANs.<sup>26</sup> Restricting the sample to the CHSDA counties also contributed to improved rate estimates, because AI/ANs in those counties are more likely to be registered for care with the IHS, and less misclassification has been noted in these counties.<sup>20</sup> Because misclassification of AI/AN race is so common in both state and national databases, <sup>18,48-51</sup> these processes contributed to improved descriptions of the lung cancer burden for AI/ANs.

Despite these improvements, there are some limitations to the data. Because most analyses were restricted to CHSDA counties, in which only 56% of AI/ANs live and which tend to be located in more rural areas and Western states (Fig. 1), the results may not be generalizable to all AI/ANs (or NHWs) in the United States. In addition, the percentage of AI/ANs residing in CHSDA counties varied greatly among the 6 IHS regions (Table 1). The percentage was particularly low in the East region (13.1%), which may help to explain the puzzling occurrence in this region of relatively low lung cancer incidence rates and high prevalence estimates of cigarette smoking.<sup>36</sup> Another limitation is that IHS records capture only enrolled members of recognized tribes who seek healthcare from the IHS and, thus, do not include others who may self-identify in the United States Census as AI/AN, notably urban AI/ANs and those who are from tribes not recognized by the federal government. Misclassification is likely to be greatest in the Pacific Coast and East regions, where the IHS has a relatively small presence and only serves a small percentage of AI/ANs.

To our knowledge, this is the most complete accounting of lung cancer occurrence among AI/AN to date and demonstrates the substantial differences in the incidence of this deadly cancer among the var-

ious AI/AN populations. Until the 1990s, most AI/ ANs were not included in areas that were covered by high-quality, population-based cancer registries. Currently, the NPCR and SEER annually produce the United States Cancer Statistics, covering 98% of United States population.<sup>21</sup> In fact, national rates of lung cancer incidence for all races combined reportedly were higher than had been noted before United States Cancer Statistics availability. 52 Although the methods used for this analysis improved the completeness of reporting cancers among AI/AN by correcting for some of the misclassification of race, the linkage protocols need to be continued. Furthermore, additional linkages with tribal rosters that were not included in the current efforts are needed to further improve the misclassification of race in AI/AN populations.

In conclusion, lung cancer is a significant problem for a large portion of AI/AN populations, notably those who live in the Northern Plains states and Alaska, where dramatic disparities in lung cancer incidence rates exist between AI/AN and NHW populations. AI/ANs in these regions have high estimates of smoking prevalence compared with NHWs<sup>36</sup> as well as high lung cancer incidence rates. The 7-fold difference between the lowest and highest lung cancer rates (Southwest vs Northern Plains) likely reflects in large part the strong association between cigarette smoking and the development of lung cancer. Indeed, BRFSS and local survey data indicate that AI/ANs in Alaska and the Plains use tobacco at rates higher than any other racial or ethnic population in the United States.<sup>39</sup> Our study underscores the need for aggressive tobacco control programs within AI/AN populations and communities.

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