

UPDATES AND EXTENSIONS TO TESTS OF THE LINEAR-NO THRESHOLD THEORY

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Two weaknesses in a 1995 article on tests of the linear-no threshold theory of radiation carcinogenesis are pointed out. One is addressed by introducing more recent cancer mortality statistics, and the other is addressed by introducing 450 newly available potential confounding factors. The later cancer statistics give results very similar to the original ones for the nation as a whole, but do cause significant changes for some geographic areas. None of the new confounding variables helps to explain the large discrepancy with predictions of the linear-no threshold theory, nor does the use of more recent cancer statistics.

Key words: Linear-no threshold theory; Radiation carcinogenesis; Confounding factors

REVIEW OF ORIGINAL STUDY

A 1995 article (1), hereafter referred to as C-95, presented a test of the linear-no threshold theory (LNT) based on a compilation of radon measurements from available sources that gave the average radon level, r , in homes for 1729 US counties, well over half of all US counties and comprising about 90% of the total US population. Plots of age-adjusted lung cancer mortality rates, m , vs. these r are shown in Figure 1a and c. To avoid the confusion of showing over a thousand data points, data have been combined into groups consisting of all counties within set intervals of r values, and for the group in each interval, the mean value of m is plotted with error bars showing 1 SD of the mean. The first and third quartiles for the distribution are also plotted as an indication of the spreads of m values involved. We see in Figure 1a and c a clear tendency for m to decrease with increasing r , in sharp contrast to the increase expected from the fact that radon can cause lung cancer, shown by the line labeled "theory."

One obvious problem is migration: people do not spend their whole lives and receive all of their radon

exposure in their county of residence at time of death where their cause of death is recorded. However, it is easy to correct the theoretical prediction for this, and the "theory" lines in Figure 1 have been so corrected. To minimize the migration problem, data for Florida, California, and Arizona were deleted because deaths there are frequently recorded for retirees who received their radon exposures elsewhere. This reduces the number of counties to 1601, but causes no appreciable change in the results.

A more serious problem is that this is an "ecological study," relating the average risk of groups (county populations) to their average exposure dose. In general, the average dose does not determine the average risk, and to assume otherwise is what epidemiologists call "the ecological fallacy." However, it is easily shown that the ecological fallacy (thus defined) does not apply in testing a linear-no threshold theory. This is familiar from the concept that, with LNT, "man-rem" determines the number of cancer deaths: dividing each by the population, man-rem gives the average dose, and number of deaths gives the mortality rate. Other problems with ecological studies that have been discussed in the epidemiology litera-

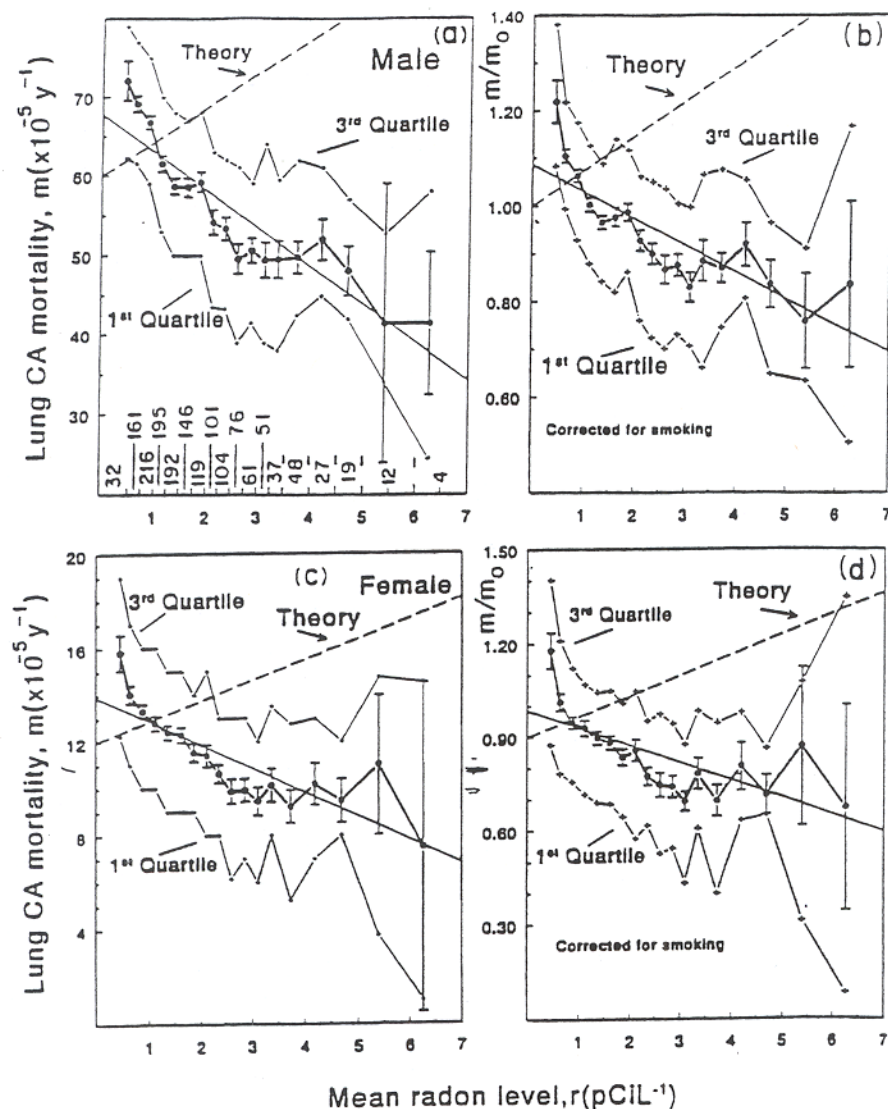


Figure 1. Lung cancer mortality rates vs. average radon levels in homes for US counties for males (a) and females (c). Data points shown are average ordinates for all counties within the range of r values shown on the baseline of (a); the number of counties within that range is also shown there. Error bars are SD of the mean, and the first and third quartiles are also shown. Lung cancer rates for (b) males and (d) females corrected for smoking prevalence (m/m_0) are shown at right. Theory lines are arbitrarily normalized lines increasing at a rate of +7.3% per pCi/L .

ture were also investigated and found not to be applicable here.

Because epidemiologists normally study the mortality risk to individuals, this study started from that premise and derived, by rigorous mathematics, the correction for smoking in terms of the fraction of adults who are smokers, S . The data in Figure 1a and c, corrected for smoking, are shown in Figure 1b and

d. It can be seen that there is a huge discrepancy between measurements and LNT theory, by about 20 SD. The theory predicts the slope of the line, B (in unit of percent per pCi/L), to be $B = +7.3$, whereas the data are fit by $B = -7.3 \pm 0.6$ and -8.3 ± 0.8 for males and females, respectively.

A wide variety of potential explanations for the discrepancy that we could develop or that were sug-

gested by others were tested and found to be grossly inadequate. Three independent sources of radon data were used but all gave the same result. Three different sources of data on smoking prevalence similarly failed. In fact, it was found that if our best estimate of the width of the distribution of S values for US counties is correct, even a perfect negative correlation between radon and smoking prevalence eliminates only half of the discrepancy. If the S value distribution had the largest credible width, an essentially perfect negative correlation between radon levels and smoking prevalence in US counties would be required to explain the discrepancy. It was shown that anything approaching such a strong correlation is completely incredible.

It was shown that the strong correlation between radon exposure and lung cancer mortality, albeit negative rather than positive, is unique to lung cancer; no remotely comparable correlation was found for any of the other 32 cancer sites. We concluded that the observed behavior is not something that can easily occur.

To investigate effects of a potential confounding variable, data were stratified into quintiles on the values of that variable, and a regression analysis was done separately for each stratum. Because the potential confounder has essentially the same value for all counties in a given stratum, its confounding effect is greatly reduced in these analyses. An average of the slopes, B , of the regression lines for the five quintiles then gives a value for B that is largely free of the confounding under investigation.

This test was carried out for 54 socioeconomic variables and none was found to be a significant confounder. In all 540 regression analyses (54 variables \times 5 quintiles \times 2 sexes), the slopes, B , were negative and the average B value for the five quintiles was always close to the value for the entire data set. This means that the negative correlation between lung cancer rates and radon exposure is found if we consider only the very urban counties or if we consider only the very rural counties; if we consider only the richest counties or only the poorest; if we consider only the counties with the best medical care or only those with the poorest medical care; and so forth for all 54 socioeconomic variables. It is also found for all strata in between, as, for example, considering only counties of average urbanicity, only counties of

average wealth, only counties of average medical care, etc.

The possibility of confounding by combinations of socioeconomic variables was studied and found not to be an important potential explanation for the discrepancy.

The stratification method was also used to investigate the possibility of confounding by geography, and of confounding by physical features such as altitude, temperature, precipitation, wind, and cloudiness, but these factors were of no help in explaining the discrepancy. The negative slope and gross discrepancy with LNT theory is found if we consider only the wettest areas, or if we consider only the driest; if we consider only the warmest areas, or only the coldest areas, etc.

The effects of the two principal recognized factors that correlate with both radon and smoking were calculated in detail: 1) urban people smoke 20% more but average 25% lower radon exposures than rural people; 2) houses of smokers have 10% lower average radon levels than houses of nonsmokers. These were found to explain only 3% of the discrepancy. Because they are typical of the largest confounding effects one can plausibly expect, it is extremely difficult to imagine a confounding effect that can explain the discrepancy. Requirements on such an unrecognized confounder were listed, and they make its existence seem extremely implausible. By far, the most plausible explanation that could be found for this discrepancy was that the linear-no threshold theory fails, grossly overestimating the cancer risk in the low dose, low dose rate region where there are no other data capable of testing the theory.

THE NEW STUDY

Two weaknesses in C-95 are addressed here by providing a great deal of further data and analysis. One weakness lies in the fact that lung cancer mortality statistics in C-95 were from 1970–1979, the latest age-adjusted rates available at that time. These deaths were presumably related to radon exposures in the 1940–1970 time period (or earlier) whereas the radon levels were measured in the 1986–1991 time period. Of course, this same problem applies to all other studies of the radon–lung cancer relation-

ship, including the widely heralded case-control studies, and it is more serious for them because it is much more likely for the radon level in a single house to have changed over several decades (e.g., cracks develop, or become sealed) than for the average radon level for all houses in a county to have changed. Nevertheless, it is preferable if the disparity in time between relevant exposures and radon measurements is reduced by using more recent lung cancer mortality statistics; that is done here. This has the added advantage of testing the degree to which these statistics have been changing with time, and it gives insight into other issues.

The other weakness addressed is the fact that an ecological study like C-95 is susceptible to "cross level bias" as emphasized by Lubin (2). For example, a case-control study may match cases and controls by annual dollar income, but an ecological study like C-95 is limited to including average annual income for different counties, not for individuals who did or did not die from lung cancer. Perhaps very poor people are much more susceptible to radon-induced lung cancer than others, making the fraction of the population that is very poor an important confounding factor. This is not necessarily represented by average income, because the latter is influenced by the fraction of the population that is very rich. The solution to this dilemma in C-95 was to consider confounding by such other factors as percent of population below the poverty line and percent unemployment.

But suppose very poor people have compensating factors (e.g., better medical care in free clinics); maybe it is people with incomes in the \$15,000–\$25,000 range that are most susceptible. The obvious solution here is to consider confounding by the fraction of the population in that income range. One might concoct models in which the fraction of the population in any other income range (or group of ranges) is an important confounder. The solution is to treat that fraction as a potential confounding factor. To do this requires data on the fraction in each income range in each county.

Of course, annual income is used here only as an example. Similar problems may arise from percent of population in any particular age range, from educational attainment, from house characteristics, etc. These problems are addressed here by considering a

very large number of potential confounding factors not available in C-95.

UPDATED LUNG CANCER RATES

The lung cancer mortality rates in C-95 were for whites only and from the time period 1970–1979; these were the latest age-adjusted rates available at that time. However, age-adjusted rates have now become available for all races from 1979 to 1994 from CDC (3), and we begin here by considering the effect of using them in the analyses. The basic equation derived and used for fitting data in C-95 was

$$m/m_o = A + B r \quad (1)$$

where

$$m_o = [S a_s + (1 - S) a_n] \quad (2)$$

and S is the fraction of the adult population that smoked cigarettes at the time their lung cancers were initiated and developed.

The determination of the constants a_n , a_s in Eq. (2) [cf. (4)] depends on the average values of m , r , and S at the relevant times, and hence must be changed to reflect changes in m when the time period is shifted. From 1970–1979 to 1979–1994, the US average m , in units of 10^{-5} /year, changed from 64.0 to 57.5 for males, and from 15.3 to 22.6 for females. This changes a_n from 8.7 to 7.8 for males, and from 3.47 to 5.13 for females, and it changes a_s from 104 to 93.5 for males and from 34.7 to 37.3 for females. With our correction for migration, the final expressions for m_o are $[8.1 + 89 S]$ for males (vs. $[9 + 99 S]$ in C-95) and $[5.3 + 48 S]$ for females (vs. $[3.7 + 32 S]$ in C-95).

In Eq. (1), m/m_o is the lung cancer rate corrected for smoking prevalence. In Figure 2, the left side shows the data for 1970–1979, reproduced from Figure 1b and d, and the right side shows the data for 1979–1994 presented in the same fashion. In accordance with Eq. (1), B is the slope of the best fit for a straight line through the data. The BEIR-IV model used in C-95 predicts B to be +7.3. In our analyses, the data from 1979–1994 mortality rates give $B = -7.7 \pm 0.51$ for males and $B = -8.2 \pm 0.67$ for females, close to the results from C-95, $B = -7.3$ and $B = -8.3$, respectively. The updated results are discrepant

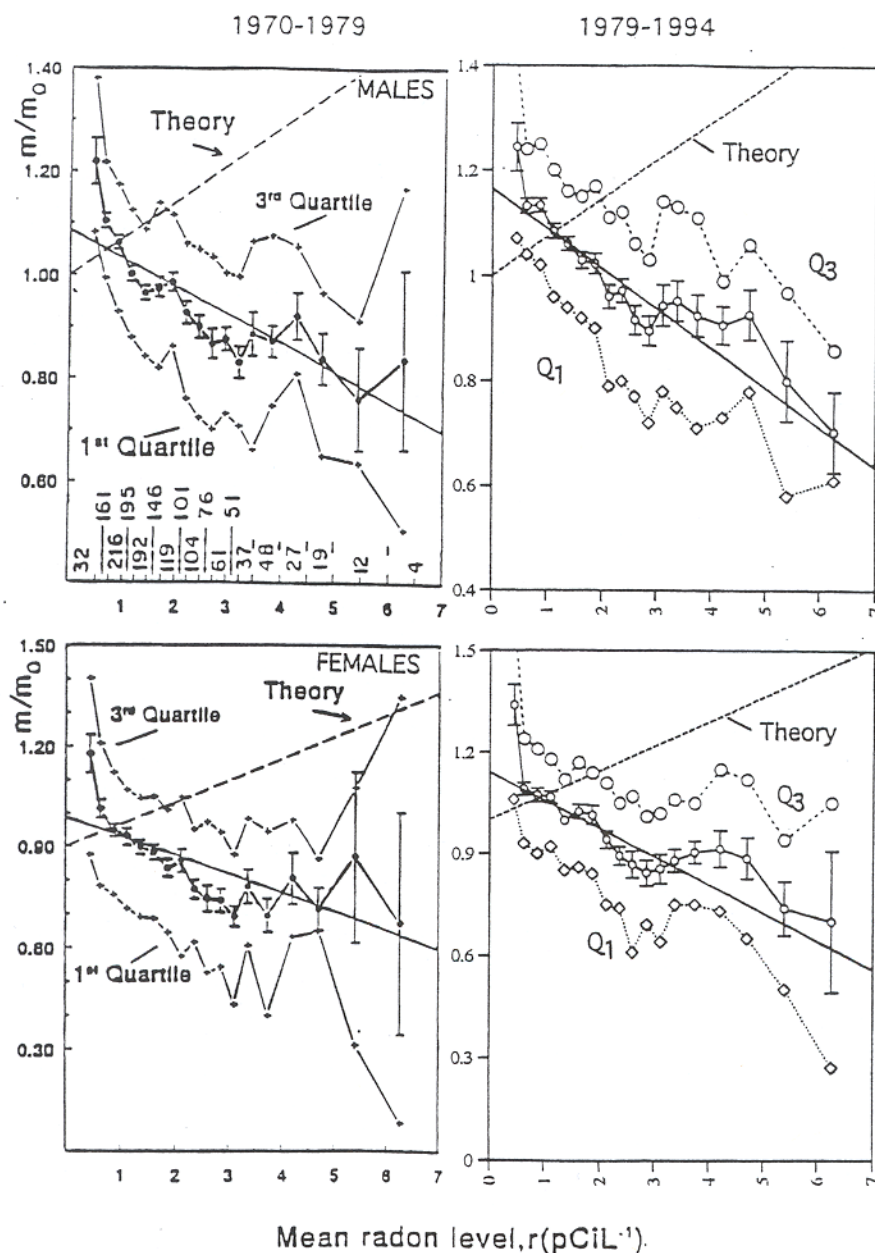


Figure 2. Comparison of results for mortality data from 1970-1979 and 1979-1994. The left side is from Figure 1b and d, lung cancer rates corrected for smoking prevalence using mortality data from 1970-1979, and the right side is a similarly treated analysis using mortality data from 1979-1994.

with the BEIR-IV prediction, $B = +7.3$, by 29 SD for males and by 23 SD for females. Further analyses will use the 1979-1994 lung cancer rates.

For some purposes, it is relevant to do similar analyses by states rather than by counties, as was done

by Cohen and Colditz (4). Because of differences in the correction for migration (moving to a different state is less common than moving to a different county), the expressions for m_0 for states are changed to $[8.0 + 88 S]$ for males and $[5.2 + 47 S]$ for females

and the BEIR-IV prediction is $B = +8.6$. Fitting the data with Eq. (1) gives $B = -11.8 \pm 2.9$ for males, and $B = -13.7 \pm 2.5$ for females, discrepant with the BEIR-IV prediction by 7.0 and 8.9 SD, respectively.

The similarity in results for the two time periods may give the impression that lung cancer rates for the two time periods are nearly identical, but that is not the case. The coefficient of correlation between these rates is 0.73 for males and 0.53 for females. For the rankings of our 1601 counties by lung cancer rates it is 0.73 and 0.61 for males and females, respectively. The average age-adjusted lung cancer rates declined by 4.5% for males and increased by 69% for females between the earlier and the later period.

ADDED SOCIOECONOMIC VARIABLES

The analyses in C-95 involved 54 socioeconomic variables (SEV) from the 1980 census. Much more extensive data have recently become available in computer transferable form from the 1990 census. About 450 of these were judged to have some possible potential for confounding the lung cancer vs. radon relationship, and these plus 23 others to be used for normalizing were transferred into our data file. As examples of normalizing factors, the number of people age >24 years who are high school graduates is normalized to the population of age >24 years, the number of people of German ancestry is normalized to the number of people reporting an ancestry, the number of people working in each occupation is normalized to the number of employed people of age >16 years, etc. As in this last example, normalization factors are sometimes not exactly appropriate, but this was judged to be unimportant in comparing different counties.

The new SEV are listed in the Appendix along with an item number, which is their column number in our data file. The 54 SEV listed in C-95 were, of course, still retained. To check on the possibility that our SEV are too finely categorized, additional SEV (items 560–577) were generated by combining several adjacent variables on annual income, value of house, ages of the population, etc. As a brief description, our personal characteristic SEV encompass age profile (31 age intervals from 0–1 years to >85 years); annual income (10 intervals from <\$5000/year to

>\$150,000/year); median and average income per capita, per family, and per household; percent below poverty level (persons, families, and households for age >65 years, with children <5 years, and with children 5–17 years); educational attainment of head of household (7 categories from <9th grade to graduate degrees). Our housing SEV include house values (20 intervals from <\$15,000 to >\$500,000), age of houses (8 intervals from <1 year to >50 years); years present occupants lived in house (6 intervals from <1 year to >30 years); housing units in structure [9 intervals from 1 to >50 (large apartment buildings)]; persons in household; age of head of household (7 intervals); number of bedrooms; 13 heating and plumbing characteristics; number of motor vehicles available; telephone availability; rent and rent as percent of income for rented houses; housing costs as percent of income for owner-occupied houses. Our ethnicity SEV include race (5 categories); ancestry (33 nationalities); languages spoken in home (17 categories); year of entry for foreign born (10 intervals); percent born in different state and different sections of US. Other categories of SEV include family type and presence and age of children, school enrollments, labor force characteristics, commuting methods and times, income type, marital status by sex and age, children ever born to women by age and by marital status, residence location 5 years previously, occupation (12 categories), industry of employment (17 categories), etc.

For a confounding factor to have an important effect on the lung cancer mortality rate (m) vs. radon (r) relationship, it must have a relatively strong correlation with m and/or r ; we designate the coefficients of correlation with these $\text{Corr-}m$ and $\text{Corr-}r$. All SEV for which either $|\text{Corr-}r|$ or $|\text{Corr-}m|$ for males is greater than 0.316 [because percent correlation = $(\text{Corr})^2$, this means correlation >10%] are listed in Table 1. Note that for every one of the 45 cases, $\text{Corr-}m$ and $\text{Corr-}r$ are of opposite sign, and only 4 of the 90 values are <0.20 (and these 4 are >0.17). For the 472 SEV under consideration, 81% of the $|\text{Corr-}r|$ and 73% of the $|\text{Corr-}m|$ are <0.20, so there is clearly a strong tendency for $|\text{Corr-}m|$ to be large when $|\text{Corr-}r|$ is large, and vice versa, and in all cases they are of opposite sign. This behavior is explained by the strong correlation between m and r seen in Figure 1. In numerical terms ($\text{Corr-}m$ for r) = ($\text{Corr-}r$ for m) = 0.40.

Table 1. Stratification on SEV With Large |Corr-*r*| or Large |Corr-*m*|

SEV No.	100 × Corr- <i>r</i>	100 × Corr- <i>m</i>	<i>B</i> Males			<i>B</i> Females		
			Q ₁	Q ₃	Q ₅	Q ₁	Q ₃	Q ₅
37	37	-37	-8.4	-8.4	-6.4	-7.0	-4.9	-7.1
82	43	-44	-8.8	-8.9	-5.3	-8.0	-6.4	-5.7
90	-34	39	-4.0	-4.4	-6.4	-7.0	-8.2	-4.1
93	20	-33	-7.9	-6.9	-7.1	-6.8	-5.0	-10.5
98	18	-34	-6.8	-5.7	-7.3	-5.7	-5.8	-7.2
111	32	-36	-7.0	-3.4	-3.5	-7.8	-7.3	-6.4
117	44	-36	-16.6	-7.2	-2.7	-13.0	-7.3	-2.6
120	-20	44	-9.6	-6.5	-1.4	-9.8	-4.9	-1.8
123	27	-40	-3.9	-5.6	-5.6	-5.1	-6.9	-9.3
132	26	-41	-5.0	-6.2	-4.2	-5.8	-9.3	-7.5
135	-24	32	-6.9	-4.8	-2.5	-8.6	-8.9	-1.8
154	37	-45	-5.5	-7.4	-2.7	-6.5	-3.7	-2.6
160	38	-46	-5.6	-5.4	-6.6	-5.4	-2.2	-8.4
178	-24	32	-8.3	-7.0	-2.6	-11.4	-6.6	-5.7
180	39	-46	-3.8	-5.4	-7.2	-3.2	-2.2	-8.7
222	33	-24	-12.6	-6.7	-7.3	-8.4	-6.3	-9.3
227	32	-31	-13.2	-2.7	-5.0	-6.8	-5.1	-10.1
228	28	-44	-8.3	-3.1	-6.5	-5.3	-5.4	-11.8
252	-17	36	-10.5	-8.9	-5.1	-12.0	-9.0	-5.7
282	44	-41	-9.0	-7.7	-5.2	-8.2	-5.0	-5.9
326	18	-34	-6.8	-5.7	-7.3	-5.7	-5.8	-7.2
342	26	-34	-3.0	-6.2	-8.2	-6.5	-7.3	-8.5
346	27	-35	-3.4	-5.2	-6.7	-4.9	-5.6	-8.6
377	32	-44	-2.9	-4.4	-7.4	+0.5	-5.5	-9.9
378	45	-48	-6.9	-7.5	-5.1	-8.1	-4.4	-4.8
388	43	-44	-7.6	-9.2	-5.3	-7.4	-6.5	-5.7
425	-32	37	-5.6	-3.2	-9.0	-5.6	-2.2	-7.5
431	-40	47	-4.7	-4.5	-8.1	-6.6	-2.8	-7.6
434	-22	41	-6.7	-6.1	7.5	-8.5	-6.4	-5.3
455	-23	36	-5.1	-7.6	-6.4	-7.5	-6.5	-8.1
459	41	-48	-8.7	-3.1	-5.7	-7.2	-5.8	-8.7
462	-39	48	-4.7	-3.5	-7.0	-7.9	-5.0	-7.4
463	-33	45	-6.5	-5.6	-10.1	-8.6	-7.5	-6.6
471	-33	42	-5.6	-5.9	-2.4	-9.5	-7.8	-3.7
490	35	-34	-13.8	-7.1	-5.3	-9.6	-8.2	-7.1
491	-36	44	-6.0	-4.4	-4.8	-6.8	-4.9	-8.2
535	26	-35	-2.5	-4.5	-7.8	-3.9	-6.8	-11.3
537	36	-46	-1.7	-6.6	-7.6	-3.3	-8.2	-9.0
538	32	-35	-5.5	-6.9	-7.2	-7.0	-6.9	-6.9
540	-28	36	-8.1	-10.4	-3.7	-8.5	-8.0	-3.3
541	-32	40	-7.3	-7.8	-2.7	-7.9	-7.7	-4.3
543	-32	33	-4.8	-6.1	-3.6	-5.8	-6.5	-5.6
549	43	-49	-3.8	-2.8	-6.2	-3.7	-2.7	-8.4
550	-32	36	-4.7	-7.7	-3.2	-5.9	-6.3	-4.7

B values for first, third, and fifth quintiles of counties from stratification on designated SEV.

To test whether an SEV is an important confounder, we stratify the data into five "quintiles" of 320 counties each, based on that SEV. In each quintile, all values of the SEV are about the same, so its confound-

ing effect is greatly reduced. Analyzing the data in each quintile independently to determine the slope, *B*, then gives values of *B* largely free of confounding by that SEV.

Results of these analyses for the first, third, and fifth quintiles, Q_1 , Q_3 , Q_5 from stratification on each SEV are listed in Table 1. For Q_3 , all values of the SEV are about average and the spread is minimal. For Q_1 , the SEV has its smallest values and for Q_5 it has its largest values. Table 1 lists results for $(45 \text{ SEV} \times 3 \text{ quintiles} \times 2 \text{ sexes} = 270 \text{ analyses})$, and in 269 of these cases, B is negative and hence grossly discrepant with the prediction of LNT theory, $B = +7.3$. The one exception, $B = +0.5$, is still below the LNT prediction by 4 SD.

Cases where B is more positive than -3.0 for any Q_1 , Q_3 , or Q_5 in Table 1 are investigated further by a finer stratification into 10 deciles (D_1 , D_2 , ..., D_{10}) of 160 counties each. For these cases, B values for D_1 and D_2 , and for D_9 and D_{10} are listed in Table 2. For SEV No. 135, 154, 180, 471, 535, 537, and 541, the finer stratification into deciles does not confirm a strong monotonic relationship between B and the SEV. For SEV No. 178, there does seem to be a trend for males, but not for females. For the three remaining cases (117—percent of German ancestry, 120—percent of Irish ancestry, 377—non-farm self-employed) there does seem to be a definite trend. For all of these, the strong correlations with radon are explainable by the urban-rural effect; people of German ancestry tend to live in rural areas, those of Irish ancestry tend to live in urban areas, and urban people are less likely to be self-employed. But this does not explain the correlation between B values and the SEV;

this correlation is not found in other SEV that reflect strong urban-rural differences.

The SEV listed in Table 2 were also investigated by multiple regression (MR). The B value (coefficient of r) in the double regression of m/m_0 on r and the SEV are listed in the columns labeled MR in Table 2. In all but one of the 22 cases, the discrepancy with the B value from LNT, $B = +7.3$, is less than the B value from single regression on r but in no case is the lessening of the discrepancy large enough to be important. In fact, even a 12 variable multiple regression of m/m_0 on r plus all 11 of the SEV listed in Table 2 gives $B = -4.1 \pm 0.50$ for males and $B = -4.5 \pm 0.73$ for females, still discrepant with the LNT prediction by 23 and 16 SD, respectively. The problem of determining B by multiple regression including numerous variables was investigated in C-95, where it was found that most of the reduction of the discrepancy between these B values and those predicted by LNT is due to methodological problems with multiple regression, and is therefore spurious.

The most important problem with multiple regression is that its relevance depends on a linear relationship. But, for the three suspicious cases, SEV No. 117, 120, and 377, a linear relationship seems quite plausible. This, plus the fact that B values listed in Table 2 for even the most extreme deciles are still much less than the LNT prediction ($+7.3$), leads one to conclude that these SEV cannot explain an appreciable part of our discrepancy.

Table 2. Finer Stratification and Multiple Regression Results for Selected SEV

SEV No.	B Males					B Females				
	D_1	D_2	D_9	D_{10}	MR	D_1	D_2	D_9	D_{10}	MR
117	-16.1	-17.5	-2.6	-2.3	-6.6	-11.0	-15.3	-1.0	-2.7	-6.4
120	-9.2	-9.1	-2.1	+1.3	-5.8	-10.0	-8.3	-2.0	-2.2	-6.4
135	-4.3	-8.6	-10.9	+3.6	-5.9	-6.5	-9.7	-6.7	+2.9	-7.9
154	-7.2	-3.8	-8.9	-2.6	-6.3	-5.3	-7.7	-8.4	-5.6	-6.0
178	-6.7	-9.3	-3.2	-1.6	-6.8	-9.5	-12.2	-8.1	-3.1	-7.7
180	-5.3	-2.2	-10.8	-3.1	-6.1	-1.6	-3.3	-11.1	-3.7	-6.0
377	-5.4	-0.3	-5.6	-8.6	-5.4	0.0	+0.9	-12.8	-7.7	-7.1
471	-1.8	-8.7	-4.6	+0.1	-5.5	-4.4	-13.8	-3.6	-3.8	-6.5
535	-3.8	-1.7	-6.7	-8.4	-6.3	-4.8	-3.3	-10.3	+1.8	-7.4
537	-8.4	+1.4	-10.8	-3.6	-5.9	-8.8	-0.4	-10.2	-6.1	-6.4
541	-6.0	-7.9	-1.7	-4.0	-6.6	-7.1	-8.2	-3.9	-4.7	-7.1

B values from analyses of first, second, ninth, and tenth deciles of counties stratified on designated SEV, and from double regression.

CONFOUNDING BY GEOGRAPHY

The one variable that is known to correlate strongly with radon levels is geography. It can be seen from Figure 1 that county average radon levels vary by an order of magnitude, and in C-95 it was found that B values obtained from analysis of separate geographic areas can vary substantially. When 1979-1994 lung cancer statistics are used, these variations were found to be somewhat enhanced.

Table 3 shows B values from the various regions and divisions of the US as defined by the Bureau of Census. We see that average B values are somewhat less negative than the -7.7 for males and -8.2 for females obtained for the nation as a whole, and the difference is increased if the geographic stratification is finer, into eight "divisions" rather than into only four "regions" (the Pacific division of the West region is not listed as it includes only WA and OR and is dominated by the former, for which results are given in Table 4).

The MINITAB statistical package we use identifies outlying data points as having large standardized residuals and extreme radon levels, which give them a strong influence on B values. The B values obtained when these are deleted are listed in the last two columns of Table 3. The one case that stands out

as important is a single county in New Hampshire (Carroll county) whose deletion changes B for females in New England from +9.4 to -0.9. In all other cases, deleting outliers has minimal effect, although always increasing the discrepancy with LNT.

In C-95, going to finer stratification on geography by considering individual states reversed the trend to less negative B with finer stratification, but that is not the case here with the 1979-1994 lung cancer rates. Data for individual states are listed in Table 4 for all states with data from more than 15 counties (for all cases listed there are 19 or more). Rather than ignore states with fewer counties, as was done in C-95, the current study combined contiguous states into groups and listed results for these groups at the end of Table 4. This allowed inclusion of all states except Delaware (3 counties) in rather natural groupings, and with no logical alternative groupings (except perhaps by rearranging the New England states). There were only 3 cases where outliers would be deleted by the prescription used in Table 3; the effects of these deletions are listed in Table 4 with asterisks.

It is interesting to ask whether the width of the distributions of B values in Table 4, as represented by their SDs (8.0 for males and 8.2 for females), can be explained simply as statistical fluctuations. To test this we take many random selections of N counties from our 1601 county data file, determine the B value

Table 3. B Values for Census Bureau Regions and Divisions

	Number Counties	Males		Females		Outliers Deleted	
		<i>B</i>	SD	<i>B</i>	SD	<i>B</i> Males	<i>B</i> Females
Regions							
Northeast	215	-2.8	1.0	-7.5	1.7	-3.4	-7.5
North Central	612	-5.0	0.8	-5.4	1.0	-5.0	-5.7
South	566	-3.9	1.2	-6.4	1.5	-5.1	-7.4
West	204	-8.9	1.4	-10.4	2.3	-9.6	-10.4
Average		-5.2		-7.4		-5.8	-7.8
Divisions							
New England	65	-0.5	4.2	+9.4	5.9	-0.5	-0.9
Mid-Atlantic	150	-3.3	1.1	-9.6	1.9	-4.0	-9.6
East North Central	308	+1.8	1.2	+1.3	1.6	+1.8	+1.3
West North Central	304	-5.7	1.1	-6.8	1.4	-5.7	-6.8
South Atlantic	273	-6.9	1.9	-7.1	2.4	-7.7	-7.1
East South Central	135	+3.2	1.5	-0.9	2.9	+2.3	-0.9
West South Central	158	-15.5	2.7	-17.4	2.8	-12.9	-17.4
Mountain	167	-5.9	1.6	-5.5	2.6	-5.9	-5.5
Average		-4.1		-4.4		-4.1	-5.9

Table 4. *B* Values for Individual States

State	Number of Counties	Males		Females	
		<i>B</i>	SD	<i>B</i>	SD
AL	38	-0.8	3.5	+1.0	5.3
AR	36	-10.9	4.2	-8.5	5.9
CO	41	-5.8	3.1	-1.7	3.6
GA	52	-1.8	5.0	-22.2	6.0
ID	40	-5.3	3.2	-8.7	5.7
IL	54	-4.1	3.1	-3.1	2.8
IN	58	-2.8	2.7	-1.4	4.3
IA	98	-1.9	1.7	-0.7	2.7
KS	21	-6.9	5.2	-7.0	5.0
KY	32	+0.8	2.6	-7.3	7.9
LA	21	-35.2	20.8	-30.4	32.3
MD	24	-13.6	5.9	-19.4	6.6
MI	53	+2.2	2.3	+2.4	3.0
MN	64	-4.6	1.4	-10.2	2.7
MS	19	+1.2	10.8	-1.6	16.2
MO	31	+1.6	8.5	+5.6	9.7
NE	43	+5.5	3.3	+5.7	4.9
		+8.5*			
NJ	21	-2.3	3.9	+3.3	5.8
NM	30	-6.3	3.8	-2.8	5.3
NY	62	+1.7	3.0	+3.5	4.6
NC	54	-5.7	3.2	-4.3	3.1
ND	38	+0.7	3.5	-0.3	5.0
OH	88	-0.2	2.0	-1.0	3.6
OK	43	-9.1	8.4	-3.8	6.8
PA	67	-1.4	1.3	-4.7	2.5
SC	36	-18.8	6.5	-1.6	12.2
TN	46	+1.7	3.0	-0.2	3.8
		-2.2*			
TX	58	-12.7	3.3	-16.9	4.0
VA	66	-6.2	4.0	-7.1	5.1
WA	29	+5.0	3.6	+6.0	8.6
WV	37	+5.4	4.6	+1.4	6.1
WI	55	-7.3	4.4	-4.5	6.2
WY	21	-7.1	6.2	+0.2	7.5
CT+MA+RI	25	-8.3	10.2	+1.6	12.8
ME+NH+VT	40	-1.0	4.9	+9.4	7.4
				-3.3*	7.7*
NV+OR+UT	32	-20.0	4.7	-10.4	10.0
MT+SD	20	-12.6	4.1	-8.5	8.5
Average		-5.0	1.3	-4.0	1.3
		-5.0*		-4.3*	
SD of distribution		8.0		8.2	

for each selection, and determine the SD of the distribution of *B* values thus determined. For example, for $N = 30$, we analyzed 60 different random selections of 30 counties; the distribution of the 60 derived *B* values had SD of 4.3 for males and 5.6 for

females. Because the number of counties in the 37 states listed in Table 4 averaged 43 and three fourths of these numbers were more than 30, we would expect the SD of the *B* values in Table 4 to be smaller than 4.3 for males and 5.6 for females if statistics

was the only consideration. The fact that this conflicts with observation means that other factors are involved. By doing calculations of the type described above for different N values, we found that the SD of about 8 found for the B values in Table 4 was what would be expected if the data for each state included only about 12 counties randomly selected from our file.

This can be understood from the fact that radon levels in the counties of a given state are not independent of one another. Because radon levels are determined by geology, and neighboring counties frequently have similar geology, the effective number of independent areas in a given state is much smaller than the number of counties. The above results suggest that a state with 43 counties typically contains about 12 truly independent areas that serve as independent data points.

This implies that the SD for all B values derived in this article should be nearly twice as large as the SD derived from our statistical treatment. For example, the B values for our entire data set, given as -7.7 ± 0.51 for males and -8.7 ± 0.67 for females, should be viewed more properly as -7.7 ± 1.0 and -8.7 ± 1.3 , respectively. This, of course, does little to resolve our discrepancy with the LNT prediction: $+7.3$.

Another aspect of Table 4 that begs for an explanation is that the average values of B (-5.0 ± 1.3 for males and -4.0 ± 1.3 for females) are quite discrepant with the values for our entire data file (-7.7 and -8.2 , respectively). Of course, all SD given for B values in this article ignore uncertainties in the data points for individual counties, so these SD are somewhat understated. But none of our sets of random selections of counties had average B values differing nearly as much from the B values for our entire 1601 county data set.

It thus seems that, for some reason we cannot explain, B values for more restricted areas tend to be less negative than for the nation as a whole. However, even these less negative B values are still grossly discrepant with the LNT prediction: $B = +7.3$.

CONCLUSION

In this article we have introduced a great deal of new data, lung cancer statistics for all races for 1979–

1994 replacing those for whites only between 1970 and 1979, and about 450 new potential confounding factors in addition to the 60+ used in C-95. The results are essentially the same as in C-95, indicating failure of the linear-no threshold theory in grossly exaggerating the risk of low-level exposure to radon.

APPENDIX: NEW SOCIOECONOMIC VARIABLES (SEV) FROM 1990 CENSUS

Item	Description
79	Total population
80	Percent urban
81	Percent rural
82	Percent living on farms
<u>School enrollment</u>	
83	Persons in schools
84	Percent of these in preprimary
85	Percent of these in elementary or high school
86	Percent of these in private schools
87	Percent of these in college
<u>Educational attainment (percent of age >25)</u>	
89	<9th grade
90	9th–12th grade, no diploma
91	High school graduates
92	Some college
93	Associates degree
94	Bachelors degree
95	Graduate or professional degree
96	High school graduates or higher
97	Bachelors degree or higher
98	Children ever born/1000 woman age 35–44
99	Percent of population born in US
100	Percent born in state of residence
101	Percent born outside US
103	Percent who speak a language other than English
104	Percent who do not speak English well
<u>Ancestry (1st or 2nd) (percent of those reporting)</u>	
106	Arab
107	Austrian
108	Belgian
109	Canadian
110	Czech
111	Danish
112	Dutch
113	English
114	Finnish
115	French
116	French Canadian
117	German
118	Greek

- 119 Hungarian
- 120 Irish
- 121 Italian
- 122 Lithuanian
- 123 Norwegian
- 124 Polish
- 125 Portuguese
- 126 Rumanian
- 127 Russian
- 128 Scotch-Irish
- 129 Scottish
- 130 Slovak
- 131 Sub-Sahara African
- 132 Swedish
- 133 Swiss
- 134 Ukrainian
- 135 American
- 136 Welsh
- 137 West Indian
- 138 Yugoslavian
- 139 Other Ancestries

Labor force

- 140 Percent in labor force
- 141 Percent of males in labor force
- 142 Percent of males unemployed
- 143 Percent of females in labor force
- 144 Percent of females unemployed

Occupation

- 146 Executive, administrative, managerial
- 147 Professional specialty
- 148 Technicians
- 149 Sales
- 150 Administrative support
- 151 Private household occupations
- 152 Protective services
- 153 Service occupations (not 151,152)
- 154 Farming, forestry, fishing
- 155 Precision production, craft, repair
- 156 Machine operators, assemblers, inspectors
- 157 Transportation
- 158 Handlers, helpers, cleaners, laborers

Industry (percent employed in)

- 160 Agriculture
- 161 Mining
- 162 Construction
- 163 Manufacturing, nondurable
- 164 Manufacturing, durable
- 165 Transportation
- 166 Communication, other utilities
- 167 Wholesale trade
- 168 Retail trade
- 169 Finance, insurance, real estate
- 170 Business services, repair services
- 171 Personal services
- 172 Entertainment, recreation
- 173 Health services

- 174 Educational services
- 175 Other professional services
- 176 Public administration

Class of worker (percent of those employed)

- 178 Private wage and salary
- 179 Government
- 180 Self-employed
- 181 Unpaid family workers

Household income (percent in each range)

- 183 <\$5000
- 184 \$5000-\$9999
- 185 \$10,000-\$14,999
- 186 \$15,000-\$24,999
- 187 \$25,000-\$34,999
- 188 \$35,000-\$49,999
- 189 \$50,000-\$74,999
- 190 \$75,000-\$99,999
- 191 \$100,000-\$149,999
- 192 >\$150,000

Annual income in dollars

- 193 Median household
- 194 Median family (Fam)
- 195 Median family (Fm)
- 196 Per capita average
- 197 Mean wage and salary
- 198 Mean non-farm self-employment
- 199 Mean farm self-employment
- 200 Mean social security
- 201 Mean public assistance
- 202 Mean retirement

Percent below poverty level

- 203 All persons
- 204 Persons age >17
- 205 Persons age >64
- 206 With related children age <5
- 207 With related children age 5-17
- 208 All families
- 209 With related children age <18
- 210 With related children age <5
- 211 Female householder families
- 212 With related children age <18
- 213 With related children age <5

Percent of housing units built in time periods

- 215 1989-1990
- 216 1985-1988
- 217 1980-1984
- 218 1970-1979
- 219 1960-1969
- 220 1950-1959
- 221 1940-1949
- 222 Before 1939

Percent with various numbers of bedrooms

- 223 No bedroom
- 224 One

- 225 Two
- 226 Three
- 227 Four
- 228 Five or more

House characteristics (percent in category)

- 229 Lacking complete plumbing facilities
- 230 Lacking complete kitchen facilities
- 231 Part of condominium
- 232 Water from public system
- 233 Water from individual drilled well
- 234 Water from individual dug well
- 235 Public sewer
- 236 Septic tank
- 237 Heated by utility gas
- 238 Heated by bottled gas, tank, or LP gas
- 239 Heated by electricity
- 240 Heated by oil
- 241 Heated by coal or coke
- 242 No heating fuel

Year householder moved into house

- 243 1989-90
- 244 1985-88
- 245 1980-84
- 246 1970-79
- 247 1960-69
- 248 Before 1960

- 249 No telephone in unit
- 251 No vehicle available
- 252 One vehicle available
- 253 Two vehicles available
- 254 Three or more vehicles available
- 255 Median mortgage (dollars)
- 256 Net median mortgage (dollars)

Owner costs as percent of household income

- 258 <20%
- 259 20%-24%
- 260 25%-29%
- 261 30%-34%
- 262 35% or more

Rent (percent of rented units in range)

- 264 <\$200
- 265 \$200-\$299
- 266 \$300-\$499
- 267 \$500-\$749
- 268 \$750-\$999
- 269 \$1000 or more
- 270 Median rent

Rent as percent of household income

- 272 <20%
- 273 20%-24%
- 274 25%-29%
- 275 30%-34%
- 276 35% or more

- 278 Percent of housing units occupied
- 279 Percent of housing units vacant
- 280 Percent urban inside urban area
- 281 Percent urban outside urban area
- 282 Percent rural—farm
- 283 Percent rural—non-farm

Units in structure (percent in category)

- 284 1, detached
- 285 1, attached
- 286 2
- 287 3 or 4
- 288 5-9
- 289 10-19
- 290 20-49
- 291 50 or more
- 292 Mobile home or trailer

Value of owner occupied units (percent in range)

- 293 <\$15,000
- 294 \$15,000-\$19,999
- 295 \$20,000-\$24,999
- 296 \$25,000-\$29,999
- 297 \$30,000-\$34,999
- 298 \$35,000-\$39,999
- 299 \$40,000-\$44,999
- 300 \$45,000-\$49,999
- 301 \$50,000-\$59,999
- 302 \$60,000-\$74,999
- 303 \$75,000-\$99,999
- 304 \$100,000-\$124,999
- 305 \$125,000-\$149,999
- 306 \$150,000-\$174,999
- 307 \$175,000-\$199,999
- 308 \$200,000-\$249,999
- 309 \$250,000-\$299,999
- 310 \$300,000-\$399,999
- 311 \$400,000-\$499,999
- 312 \$500,000 or more
- 313 Median value
- 314 Median value with mortgage
- 315 Median value not mortgaged

Residence five years earlier (percent in category)

- 317 Same house
- 318 Different house in US
- 319 Different house in same state
- 320 Different house in same county
- 321 House in different county, same state
- 322 House in different state
- 323 Lived abroad

Children ever born per 1000 women

- 324 Women age 15-24 years
- 325 Women age 25-34 years
- 326 Women age 35-44 years
- 328 Percent of foreign born entered US 1980-1990

- 330 Percent of population speaking other than English
- 331 Percent not speaking English "very well"
- 332 Percent speaking Spanish
- 333 Percent speaking Spanish, not English "very well"
- 334 Percent speaking Asian language
- 335 Percent speaking Asian, not English "very well"
- 336 Civilian labor force
- 337 Percent of civilian labor force employed
- 338 Percent of civilian labor force unemployed
- 339 Percent of civilian labor force unemployed
- 340 Percent of labor force in armed forces

Commuting to work

- 342 Percent drive alone
- 343 Percent in carpools
- 344 Percent using public transportation
- 345 Percent using other means
- 346 Percent walk or work at home
- 347 Mean travel time (minutes)
- 349 Percent of employment by government
- 350 Percent employment by local government
- 351 Percent employment by state government
- 352 Percent employment by federal government

Percent of family households with income of

- 354 <\$5000
- 355 \$5000-\$9999
- 356 \$10,000-\$14,999
- 357 \$15,000-\$24,999
- 358 \$25,000-\$34,999
- 359 \$35,000-\$49,999
- 360 \$50,000-\$74,999
- 361 \$75,000-\$99,999
- 362 \$100,000-\$149,999
- 363 >\$150,000

Percent of nonfamily households with income of

- 365 <\$5000
- 366 \$5000-\$9999
- 367 \$10,000-\$14,999
- 368 \$15,000-\$24,999
- 369 \$25,000-\$34,999
- 370 \$35,000-\$49,999
- 371 \$50,000-\$74,999
- 372 \$75,000-\$99,999
- 373 \$100,000-\$149,999
- 374 >\$150,000

Percent of households with income from

- 376 Wage & salary
- 377 Non-farm self-employment
- 378 Farm self-employment
- 379 Social Security
- 380 Public assistance
- 381 Retirement plan
- 386 Percent of persons urban, inside urbanized area
- 387 Percent of persons urban, outside urbanized area
- 388 Percent of persons rural, farm

- 389 Percent of persons rural, non-farm
- 390 Percent of persons male
- 391 Percent of persons female

Percent of persons in age range

- 392 <1 year
- 393 1-2 years
- 394 3-4 years
- 395 5 years
- 396 6 years
- 397 7-9 years
- 398 10-11 years
- 399 12-13 years
- 400 14 years
- 401 15 years
- 402 16 years
- 403 17 years
- 404 18 years
- 405 19 years
- 406 20 years
- 407 21 years
- 408 22-24 years
- 409 25-29 years
- 410 30-34 years
- 411 35-39 years
- 412 40-44 years
- 413 45-49 years
- 414 50-54 years
- 415 55-59 years
- 416 60-61 years
- 417 62-64 years
- 418 65-69 years
- 419 70-74 years
- 420 75-79 years
- 421 80-84 years
- 422 85 years and over

Percent of persons age >14 in category

- 423 Male, never married
- 424 Male, married, spouse present
- 425 Male, married, spouse absent, separated
- 426 Male, married, spouse absent, other reasons
- 427 Male, widowed
- 428 Male, divorced
- 429 Female, never married
- 430 Female, married, spouse present
- 431 Female, spouse absent, separated
- 432 Female, spouse absent, other reasons
- 433 Female, widowed
- 434 Female, divorced
- 435 Female, never married, age 15-24
- 436 Female, never married, age 25-34
- 437 Female, never married, age 35-44
- 438 Female, never married, age 45 and over
- 439 Female, ever married, age 15-24
- 440 Female, ever married, age 25-34
- 441 Female, ever married, age 35-44
- 442 Female, ever married, age 45 and over

Percent of persons in group quarters

- 443 Correctional institutions
- 444 Nursing homes
- 445 Mental hospitals
- 446 Juvenile institutions
- 447 Other institutions
- 448 College dormitories
- 449 Military quarters
- 450 Shelters for homeless
- 451 Visible in street locations

Percent of persons born in

- 452 State of residence
- 453 Other state in Northeast
- 454 Other state in Midwest
- 455 Other state in South
- 456 Other state in West

Percent of families with

- 458 Married couple, children >17 years
- 459 Married couple, no children >17 years
- 460 Male householder, no wife, children >17 years
- 461 Male householder, no wife, no children >17 years
- 462 Female householder, no husband, children >17 years
- 463 Female householder, no husband, no children >17 years
- 464 No workers
- 465 1 worker
- 466 2 workers
- 467 3 or more workers

Percent of households with

- 469 1 person
- 470 2 persons
- 471 3 persons
- 472 4 persons
- 473 5 persons
- 474 6 persons
- 475 7 persons or more

Percent of family households with householder age

- 476 15-24 years
- 477 25-34 years
- 478 35-44 years
- 479 45-54 years
- 480 55-64 years
- 481 65-74 years
- 482 75 years and over

Percent of nonfamily households with householder age

- 483 15-24 years
- 484 25-34 years
- 485 35-44 years
- 486 45-54 years
- 487 55-64 years
- 488 65-74 years
- 489 75 years and over

Percent of persons

- 490 White
- 491 Black

- 492 American Indian or Aleut

- 493 Asian or Pacific Islander

- 494 Other race

- 495 Hispanic origin, white

- 496 Hispanic origin, black

- 497 Hispanic origin, American Indian

- 498 Hispanic origin, Asian or Pacific Islander

- 499 Hispanic origin, other race

Percent of foreign-born persons entered US in

- 500 1987-1990

- 501 1985-1986

- 502 1982-1984

- 503 1980-1981

- 504 1975-1979

- 505 1970-1974

- 506 1965-1969

- 507 1960-1964

- 508 1950-1959

- 509 Before 1950

Language spoken in home

- 510 Only English

- 511 German

- 512 Yiddish

- 513 Other West Germanic

- 514 Scandinavian

- 515 Greek

- 516 Italian

- 517 French

- 518 Spanish

- 519 Polish

- 520 Russian

- 521 Chinese

- 522 Hungarian

- 523 Japanese

- 524 Korean

- 525 North American Indian

- 526 Vietnamese

Transportation to work

- 527 Drive alone

- 528 Carpool

- 529 Bus or trolley bus

- 530 Street car or trolley car

- 531 Subway or elevated

- 532 Railroad

- 533 Motorcycle

- 534 Bicycle

- 535 Walk

- 536 Other

Travel time to work (minutes)

- 537 Less than 5

- 538 5-9

- 539 10-14

- 540 15-19

- 541 20-24

- 542 25-29

543 30-34
 544 35-39
 545 40-44
 546 45-59
 547 60-89
 548 90 or more
 549 Work at home
 550 Total

COMBINED CATEGORIES

560 Household income <\$25,000
 561 Household income \$25,000-\$75,000
 562 Household income >\$75,000
 563 Housing built after 1979
 564 Housing built 1950-1979
 565 Housing built before 1950
 566 10 or more units in structure

Owner-occupied housing unit value

567 <\$40,000
 568 \$40,000-\$100,000
 569 \$100,000-\$250,000
 570 >\$250,000

Percent of population in age range

571 0-17 years

572 18-34 years
 573 35-64 years
 574 65 years or more

Percent of foreign born entered US

575 Since 1979
 576 1965-1979
 577 Before 1965

REFERENCES

1. Cohen, B. L. Test of the linear-no threshold theory of radiation carcinogenesis for inhaled radon decay products. *Health Phys.* 68:157-174; 1995.
2. Lubin, J. H. On the discrepancy between ecologic studies in individuals of lung cancer and residential radon and Cohen's ecologic regression. *Health Phys.* 75:4-10; 1998.
3. Centers for Disease Control and Prevention. Wonder searches and queries; 1997. <http://wonder.cdc.gov/rchtml/Convert/data/AdHoc.html>
4. Cohen, B. L.; Colditz, G. A. Test of the linear-no threshold theory of radiation carcinogenesis. *Environ. Res.* 64:65-89; 1994.