

## PAPERS AND SHORT REPORTS

## Non-smoking wives of heavy smokers have a higher risk of lung cancer: a study from Japan

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### Abstract

In a study in 29 health centre districts in Japan 91 540 non-smoking wives aged 40 and above were followed up for 14 years (1966-79), and standardised mortality rates for lung cancer were assessed according to the smoking habits of their husbands. Wives of heavy smokers were found to have a higher risk of developing lung cancer and a dose-response relation was observed. The relation between the husband's smoking and the wife's risk of developing lung cancer showed a similar pattern when analysed by age and occupation of the husband. The risk was particularly great in agricultural families when the husbands were aged 40-59 at enrolment. The husbands' smoking habit did not affect their wives' risk of dying from other disease such as stomach cancer, cervical cancer, and ischaemic heart disease. The risk of developing emphysema and asthma seemed to be higher in non-smoking wives of heavy smokers but the effect was not statistically significant.

The husband's drinking habit seemed to have no effect on any causes of death in their wives, including lung cancer.

These results indicate the possible importance of passive or indirect smoking as one of the causal factors of lung cancer. They also appear to explain the long-standing riddle of why many women develop lung cancer although they themselves are non-smokers. These results also cast doubt on the practice of assessing the relative risk of developing lung cancer in smokers by comparing them with non-smokers.

### Introduction

The possible consequences to the health of non-smokers of long-term exposure to cigarette smoke (passive smoking) should

be studied thoroughly because the side-stream and second-hand smoke of cigarettes contain various toxic substances, including carcinogens.<sup>1 2</sup> The need for such a study increased by the report of small-airways dysfunction in non-smokers chronically exposed to tobacco smoke.<sup>3</sup>

The effect of passive smoking on lung cancer was studied by following 91 540 non-smoking housewives aged 40 and above and measuring their risk of developing lung cancer according to the smoking habits of their husbands.

### Methods

To study the consequences to health of such factors as cigarette smoking, alcohol drinking, occupation, and marital status, a prospective population study has been in progress in 29 health centre districts in six prefectures in Japan since the autumn of 1965. In total 265 118 adults (122 261 men and 142 857 women) aged 40 years and over, 91-99% of the census population, were interviewed and followed by establishing a record linkage system between the risk-factor records, a residence list obtained by special yearly census, and death certificates.

Since the effect of direct smoking of cigarettes in this study has already been reported,<sup>4-7</sup> my study focused on the effect of husband's smoking on the risk of lung cancer in their non-smoking wives. Such observation was possible since detailed questions about lifestyle, including smoking habits, were asked of husbands and wives independently at the start of this study. No subjective bias was therefore conceivable.

A total of 346 deaths from lung cancer in women were recorded during 14 years of follow-up (1966-79). Of these women 245 were married, and 174 of these were also non-smokers. These cases occurred among 91 540 non-smoking married women whose husbands' smoking habits were studied. The risk of lung cancer was carefully measured, taking into consideration possible confounding variables.

### Results

Wives of heavy smokers were found to have a higher risk of developing lung cancer than wives of non-smokers and a statistically significant dose-response relationship was observed (Mantel-extension  $\chi$  test result being 3.299; two-tailed  $p=0.00097$ ). Age-occupation standardised annual mortality rates for lung cancer were 8.7/100 000 (32 out of 21 895) when husbands were non-smokers or occasional smokers,

14.0 (86 out of 44 184) when husbands were ex-smokers or daily smokers of 1-19 cigarettes, and 18.1 (56 out of 25 146) when husbands were daily smokers of 20 or more cigarettes. These figures gave risk ratios of 1.00, 1.61, and 2.08 respectively. A similar trend was observed in age and occupation groups of husbands (table I).

TABLE I—Standardised mortality for lung cancer in women by age, occupation, and smoking habit of the husband (patient herself a non-smoker)

Husband's smoking habit:	Non-smoker	Ex-smoker or 1-19/day	≥20/day
<i>Husband's age: 40-59 years</i>			
Population of wives	14 020	30 676	20 584
No of deaths from lung cancer	11	40	36
Occupation-standardised mortality/100 000	5.64	9.34	13.14
<i>Husbands age: ≥60 years</i>			
Population of wives	7875	13 508	4877
No of deaths from lung cancer	21	46	20
Occupation-standardised mortality/100 000	15.79	24.44	29.60
Standardised risk ratio for all ages	1.00	1.61	2.08
<i>Husband working in agriculture</i>			
Population of wives	10 406	20 044	9391
No of deaths from lung cancer	17	52	24
Age-standardised mortality/100 000	9.54	17.02	18.40
<i>Husband working elsewhere</i>			
Population of wives	11 489	24 140	16 070
No of deaths from lung cancer	15	34	32
Age-standardised mortality/100 000	9.13	10.46	17.78
Standardised risk ratio for all occupations	1.00	1.43	1.90

The relation between the husband's smoking habit and the wife's risk of developing lung cancer was particularly significant in agricultural families when the husband was aged 40-59 at enrolment (Mantel-extension chi being 2.597 or two-tailed  $p=0.0094$ ); lung cancer risk ratios were 1.00, 3.17, and 4.57 when husbands were non-smokers or occasional smokers, ex-smokers or smokers of 1-19 cigarettes daily, and smokers of 20 or more cigarettes daily respectively (table II).

TABLE II—Mortality for lung cancer in women by occupation and by smoking habit of husband among men aged 40-59 (patient herself a non-smoker)

Husband's smoking habit:	Non-smoker	Ex-smoker or 1-19/day	≥20/day
<i>Agricultural workers:</i>			
Population of wives	5 999	12 753	7150
No of deaths from lung cancer	3	20	16
Mortality/100 000	3.48	11.03	15.92
<i>Other workers</i>			
Population of wives	8 021	17 923	13 434
No of deaths from lung cancer	8	20	20
Mortality/100 000	7.15	8.09	11.05
Standardised risk ratio for all occupations	1.00	1.67	2.36

The husbands' smoking habits seemed to have no effect on their wives' risk of developing other major cancers, such as cancers of the stomach ( $n=716$ ) and of the cervix ( $n=250$ ) or ischaemic heart disease ( $n=406$ ). The risk of developing emphysema and asthma seemed to be higher among the non-smoking wives of smokers, but the effect was not statistically significant (table III).

Other characteristics of the husbands, such as their alcohol drinking habits did not affect mortality from lung cancer in their wives. The relative risk ratios of death from lung cancer were 1.00, 1.13, and 1.18 ( $p=0.396$ ) respectively when husbands were non-drinkers, occasional or rare drinkers, and daily drinkers. Similar results were found with other causes of death (table IV).

Finally, the effect of passive smoking was compared with the effect of direct smoking. The effect of passive smoking was around one-half to one-third that of direct smoking. The relative risk of developing lung cancer by passive smoking was about 1.8 compared with about 3.8 in direct smokers (fig 1).

TABLE III—Age-occupation standardised risk ratio for selected causes of death in women by smoking habit of the husband (patient herself a non-smoker)

Cause of death	Husband's smoking habit			p value
	Non-smoker	Ex-smoker, or 1-19/day	≥20/day	
Lung cancer ( $n=174$ )	1.00	1.61	2.08	0.001
Emphysema, asthma ( $n=66$ )	1.00	1.29	1.49	0.474
Cancer of cervix ( $n=250$ )	1.00	1.15	1.14	0.249
Stomach cancer ( $n=716$ )	1.00	1.02	0.99	0.720
Ischaemic heart disease ( $n=406$ )	1.00	0.97	1.03	0.393

TABLE IV—Age-standardised risk ratio for selected causes of death in women by alcohol-drinking habit of the husband

Cause of death	Husband's drinking habit			p value
	Non-drinker	Occasional or rare drinker	Daily drinker	
Lung cancer ( $n=174$ )	1.00	1.13	1.18	0.396
Emphysema, asthma ( $n=66$ )	1.00	0.92	1.39	0.292
Cancer of cervix ( $n=250$ )	1.00	0.84	0.89	0.514
Stomach cancer ( $n=716$ )	1.00	0.88	0.95	0.285
Ischaemic heart disease ( $n=406$ )	1.00	1.09	0.93	0.567

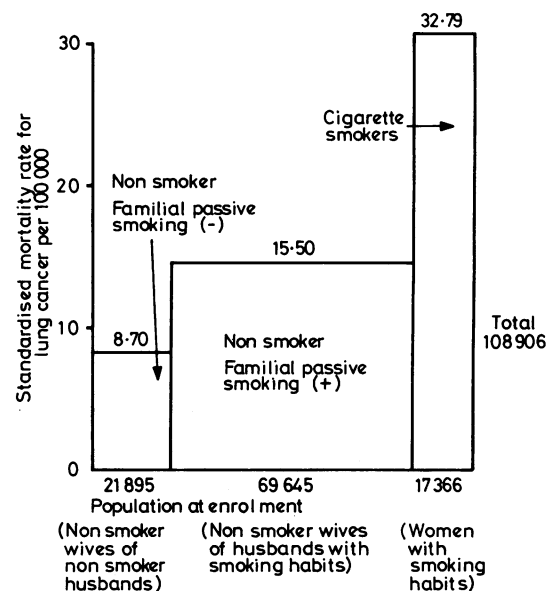


FIG 1—Lung cancer mortality in women according to the presence or absence of direct and familial indirect smoking.

## Discussion

The possible effect of passive smoking was studied by following many non-smoking wives whose husbands had various smoking habits, and measuring their risk of developing lung cancer. Continued exposure to their husbands' smoking increased mortality from lung cancer in non-smokers up to twofold. The extent of the increase in the risk of developing cancer reached as high as 4.6 for non-smoking wives of agricultural workers aged 40-59 who smoked 20 or more cigarettes a day.

The fact that there was a statistically significant relation (two-tailed  $p=0.00097$ ) between the amount the husbands smoked and the mortality of their non-smoking wives from lung cancer suggests that these findings were not the result of chance. To determine whether such an effect was limited to lung cancer, similar studies were conducted with other causes of death. Although there seemed to be a relation between husbands' smoking habits and deaths from emphysema and asthma in their wives, the effect of passive smoking was strongest with

lung cancer. Passive smoking did not seem to increase the risk of developing stomach cancer, cervical cancer, or ischaemic heart disease. We found that smoking was the only habit of the husbands to affect wives' mortality. The absence of an effect of husbands' drinking habits on mortality in their wives was shown as an example.

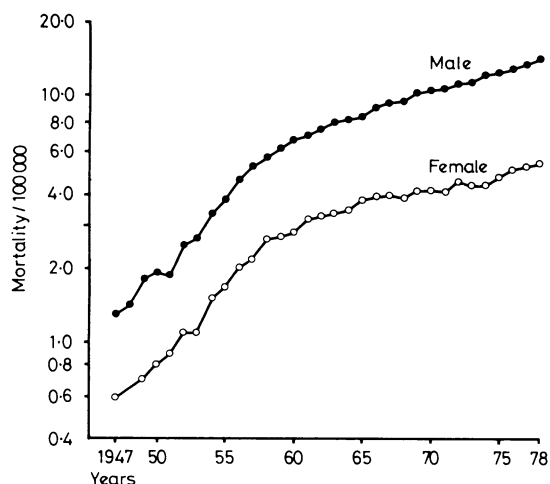


FIG 2—Age-adjusted mortality for lung cancer in Japan (1947-78).

The most important confounding variables would have been urban factors. Similar observations were therefore made for agricultural families and for non-agricultural families, and a similar dose-response relation was observed in both groups. The effect of passive smoking was most striking in younger couples in agricultural families, relative risk reaching 4-6, probably because of the lesser extent of the exposure to passive smoking outside the family in the case of rural residents. That the rate for non-smoking wives with husbands who were heavy smokers in urban families was lower than that in rural families is puzzling but probably reflects a longer period of mutual contact of couples in rural families. In urban families some couples meet only for a short period in the day.

Finally, the effects of passive smoking were compared with the effects direct smoking. The results clearly indicated that the effect of passive smoking is about one-half to one-third that of direct smoking in terms of mortality ratio or relative risk. In terms of attributable risk, however, the effect of passive smoking on lung cancer in women must be much more important than that of direct smoking (fig 1), especially in countries such as

Japan where 73% of men but only 15% of women smoke. Therefore, although the relative risk of indirect smoking was smaller than that of direct smoking, the absolute excess deaths from lung cancer due to passive smoking must be important because of the large size of the exposed group.

The age-adjusted mortality rates for lung cancer have been sharply increasing both for men and for women in Japan (fig 2). As only a fraction of Japanese women with lung cancer smoke cigarettes, the reasons why their mortality from lung cancer parallels that in men have been unclear. The present study appears to explain at least a part of this long-standing riddle.

This observation also questions the validity of the conventional method of assessing the relative risk of developing lung cancer in smokers by comparing them with non-smokers. This study shows that non-smokers are not a homogenous group and should be subdivided according to the extent of previous exposure to indirect or passive smoking.

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ARSSMART. The hot Arssmart is called also Water-pepper, or Culrage. The mild Arssmart is called dead Arssmart Persicaria, or Peachwort, because the leaves are so like the leaves of a peach-tree; it is also called Plumbago.

The mild has broad leaves set at the great red joint of the stalks; with semicircular blackish marks on them, usually either blueish or whitish, with such like seed following. The root is long, with many strings thereat, perishing yearly; this has no sharp taste (as another sort has, which is quick and biting) but rather sour like sorrel, or else a little drying, or without taste. It grows in watery places, ditches, and the like, which for the most part are dry in summer. It flowers in June, and the seed is ripe in August.

As the virtue of both these is various, so is also their government; for that which is hot and biting, is under the dominion of Mars, but Saturn, challenges the other, as appears by that leaden coloured spot he hath placed upon the leaf.

It is of a cooling and drying quality and very effectual for putrified ulcers in man or beast, to kill worms, and cleanse the putrified places. The juice thereof dropped in, or otherwise applied, consumes all colds, swellings, and dissolveth the congealed blood of bruises by strokes, falls, etc. A piece of the root, or some of the seeds bruised, and

held to an aching tooth, takes away the pain. The leaves bruised and laid to the joint that has a felon thereon, takes it away. The juice destroys worms in the ears, being dropped into them; if the hot Arssmart be strewed in a chamber, it will soon kill all the fleas; and the herb or juice of the cold Arssmart, put to a horse or other cattle's sores, will drive away the fly in the hottest time of Summer; a good handful of the hot biting Arssmart put under a horse's saddle, will make him travel the better, although he were half tired before. The mild Arssmart is good against all imposthumes and inflammations at the beginning, and to heal green wounds.

All authors chop the virtues of both sorts of Arssmart together, as men chop herbs for the pot, when both of them are of contrary qualities. The hot Arssmart grows not so high or tall as the mild doth, but has many leaves of the colour of peach leaves, very seldom or never spotted; in other particulars it is like the former, but may easily be known from it, if you will but be pleased to break a leaf of it cross your tongue, for the hot will make your tongue to smart, but the cold will not. If you see them both together, you may easily distinguish them, because the mild hath far broader leaves. (Nicholas Culpeper (1616-54) *The Complete Herbal*, 1850.)