Mortality from alcohol related disease in Italy

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SUMMARY Trends in death certification rates from the five major alcohol related causes of death in Italy (cancers of the mouth or pharynx, oesophagus, larynx, liver and cirrhosis of the liver) were analysed over a period (1955-79) in which *per capita* alcohol consumption almost trebled. Age standardised mortality from liver cirrhosis almost doubled in males and increased over 70% in females. In males, mortality from cancers of the upper digestive or respiratory tract showed increases of between 27% and 44%, and liver cancer increased by over 100%. In the late 1970s, the four alcohol related cancer sites accounted for about 12% of all cancer deaths in males and 4.5% in females. Mortality from liver cirrhosis alone accounted for 4.8% of all deaths in males (9.2% of manpower years lost) and 2.3% in females (6.3% manpower years lost) in females. These figures were even higher in selected areas of north eastern Italy, where alcohol consumption is greater. In absolute terms, the upward trends observed correspond to about 10 000 excess deaths per year in the late 1970s compared with rates observed two decades earlier and are thus second only to the increase in tobacco related causes of death over the same calendar period.

Although it is known that alcohol consumption in Italy has been traditionally high,¹ the extent to which it has risen over the last few decades is probably less appreciated: between 1950 and 1980, in fact, national *per capita* consumption values rose over threefold.^{2,3} A major difficulty in assessing the health consequences of changes in alcohol consumption in Italy arises from the pattern of drinking, which chiefly involves long term heavy drinking, "binges" being relatively infrequent. Not surprisingly, therefore, only about 350 deaths per year in the late 1970s were certified as due to "alcoholism" and 50 to "alcoholic psychosis".⁴

Diseases and deaths related to constant heavy drinking each day are in fact much more frequent. In the late 1970s over 20 000 death certifications per year were due to liver cirrhosis and over 10 000 to the alcohol related cancer sites (mouth or pharynx, oesophagus, larynx, and liver),⁵ out of a total of about 500 000 deaths per year in the whole of Italy.⁴

In the present study, national trends in mortality rates from the major alcohol related causes of death over the period 1955-79 are presented, and changes in rates within different geographical areas are discussed.

Materials and methods

Death certification numbers in the whole of Italy for the causes of interest, stratified for sex and age in five-year groups, were abstracted from publications by the Italian Central Institute of Statistics (ISTAT), available from 1955 onwards.⁶ There were no major changes in the definition or coding of the causes of death considered between the Seventh, Eighth and Ninth Revisions of the International Classification of Diseases (ICD, table 1). Estimates of the resident population were obtained from publications by the Department of Demography of the University of

Table 1 Codes of the International Classification of Diseases (ICD) included in each category of neoplasms considered

Type of cancer	7th ICD	8th ICD	9th ICD
Mouth or pharynx	140-148	140-149	140-149
Oesophagus	150	150	150
Larynx	161	161	161
Liver	155.0 + 155.8	155	155.0

Rome (for the period 1955–72),⁷ and by the Central Institute of Statistics from 1972 onwards.⁸ On the basis of these data, age specific and age standardised mortality rates were computed, age standardisation being performed by the direct method using the 1971 census population (subdivided in five year age groups) as standard.

Copies of original computer tapes were also obtained from the ISTAT with extracts of all primary death records for the period 1975–77, since on the basis of published material death certification numbers are not available stratified simultaneously for age, sex, and province. From these data age specific directly standardised mortality rates and standardised mortality ratios from the causes of interest in each of the 95 Italian provinces were computed.

Results

Age standardised, truncated 35–64 years death certification rates from the four alcohol related cancer sites and liver cirrhosis in the whole of Italy over the period 1955–79 are reported in table 2. In males,

mortality from cancers of the mouth or pharynx, oesophagus and larynx showed increases of between 27% and 44%, and mortality from primary cancer or cirrhosis of the liver roughly doubled. In females, death certification rates from cancers of the mouth or pharynx, larynx or oesophagus were appreciably lower than in males and showed no consistent trend. There was a moderate upward trend in mortality from cancer of the liver and an over 70% increase for liver cirrhosis.

In the late 1970s, the four alcohol related cancer sites accounted for about 12% of all cancer deaths in males and 4.5% in females (overall, 8.8%). Mortality from liver cirrhosis alone accounted for 4.8% of all deaths in males and 2.3% in females. These proportions rose to 9.2% for males and 6.3% for females when manpower years lost (between the ages of 25 and 65) were computed. Overall the five alcohol related causes considered accounted for about 6% of all deaths.

When age-specific death certification rates from liver cirrhosis were considered (table 3), the upward trends were much more marked in the younger age

Table 2 Age standardised Italian death certification rates /100 000 people aged 35–64 from selected alcohol-related diseases

Cause of death	Sex	Rates/100 000 people aged 35–64 for:					Rate of change 1955–79	
		1955–59	1960–64	1965–69	1970–74	1975–79	Absolute change	Percent change /yr*
Cancer of mouth or pharynx	M	6.78	7.18	7.34	7.54	7.53	+ 0.75	+ 0.5
	F	1.50	1.55	1.53	1.47	1.38	– 0.12	– 0.4
Cancer of oesophagus	M	5.27	5.71	5.86	5.89	6.13	+ 0.86	+ 0.8
	F	1.43	1.47	1.40	1.38	1.43	–	–
Cancer of larynx	M	5.23	6.21	7.38	8.23	8.48	+ 3.25	+ 2.4
	F	0.45	0.52	0.48	0.51	0.54	+ 0.09	+ 0.9
Cancer of liver	M	2.40	3.10	3.47	4.09	5.37	+ 2.97	+ 4.0
	F	2.33	2.70	2.92	2.95	3.61	+ 1.28	+ 2.2
Liver cirrhosis	M	25.00	31.37	39.07	45.16	47.36	+ 22.36	+ 3.2
	F	10.17	12.11	14.79	17.18	18.56	+ 8.39	+ 3.0

*Average annual rate of change, assuming that the change has been linear and approximated as $100\% \times \text{difference in } \log_e \text{ rates divided by 20 years}$.

Table 3 Age specific Italian death certification rates/100 000 people of selected age groups from liver cirrhosis

Sex	Age group	Rates/100 000 people for:					Rate of change 1955–79	
		1955–59	1960–64	1965–69	1970–74	1975–79	Absolute change	Percent change /yr*
Males	35–44	8.73	12.86	19.41	24.51	24.63	+ 15.90	+ 5.2
	45–54	33.16	42.11	54.09	68.11	74.18	+ 41.02	+ 4.0
	55–64	83.23	98.93	119.73	133.00	140.86	+ 57.63	+ 2.6
	65–74	125.12	157.63	185.20	202.76	208.50	+ 83.38	+ 2.6
Females	35–44	3.14	4.21	6.06	7.20	8.06	+ 4.92	+ 4.7
	45–54	11.58	13.28	16.99	19.47	20.73	+ 9.15	+ 2.9
	55–64	27.11	31.84	36.40	40.03	43.90	+ 16.79	+ 2.4
	65–74	39.73	48.05	57.84	65.16	70.42	+ 30.69	+ 2.9

*Average annual rate of change, assuming that the change has been linear and approximated as $100\% \times \text{difference in } \log_e \text{ rates divided by 20 years}$.

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Table 4 Age specific Italian death certification rates/100 000 males of selected age groups from cancers of the mouth or pharynx and oesophagus

Cause of death	Age group	Rates/100 000 males for:					Rate of change 1955-79	
		1955-59	1960-64	1965-69	1970-79	1975-79	Absolute change	Percent change/yr*
Cancer of mouth or pharynx	35-44	1.11	1.38	1.93	2.14	2.32	+1.21	+3.7
	45-54	5.52	6.07	7.05	8.74	11.06	+5.54	+3.5
	55-64	17.71	18.91	19.27	19.74	22.26	+4.55	+1.1
	65-74	35.52	36.88	36.66	34.67	32.63	-2.89	-0.4
Cancer of oesophagus	35-44	0.60	0.42	0.93	1.00	1.10	+0.50	+3.0
	45-54	3.90	4.04	4.18	5.01	6.71	+2.81	+2.7
	55-64	14.86	15.38	15.50	16.10	17.26	+2.40	+0.7
	65-74	31.41	35.06	33.07	31.91	29.92	-1.49	-0.2

*Average annual rate of change, assuming that the change has been linear and approximated as $100\% \times \text{difference in } \log_e \text{ rates divided by 20 years}$.

groups for both sexes. Steady rises, however, were also evident in middle and later age.

The pattern of age specific trends is different for cancers of the mouth or pharynx and oesophagus in males. As shown in table 4, in fact, large increases were observed in younger males, but mortality rates were roughly constant (or slightly downwards) in older age groups. When mortality rates were plotted against the central year of birth cohort⁹ (data not shown), decreases were evident for cohorts born between 1900 and 1915, followed by marked upward trends in subsequent cohorts. Quite interestingly in terms of alcohol related carcinogenesis, there was a marked fall (over 70%) in *per capita* alcohol consumption (derived from trade and sale data² in Italy between 1920 and 1935) followed by steady and marked increases later (fig. 1).

The geographical distribution of mortality from liver cirrhosis in various Italian provinces is shown in figure 2. Markedly elevated ratios in both sexes were observed in the north, chiefly in north eastern areas where alcohol consumption is greater. For instance, in the early 1970s, *per capita* wine consumption estimated within a national household survey¹⁰ was 143, and 108 litres per year in the two major north eastern regions (Veneto and Friuli Venezia Giulia) as compared with a national average of 87 litres. For females, a few southern provinces showed elevated cirrhosis mortality ratios. This suggested, however indirectly, that causes other than alcohol consumption (ie, hepatitis B virus) may have a proportionally greater importance for females.

Analyses of the four alcohol related cancer sites, reported in a separate publication,¹¹ produced largely comparable patterns for males (though somewhat less clearly for liver cancer). Consequently, for males significant positive correlations were evident between mortality rates from these causes in various Italian provinces, ranging from +0.38 between cancer of the larynx and liver cirrhosis and +0.71 between cancers of the oesophagus and larynx.

Fig 1 Trends in per capita total alcohol consumption in Italy (averaged to unity over the whole period).

Discussion

The findings of this analysis of trends in death certification rates from alcohol related diseases in Italy are extremely worrying, since overall mortality from the five major alcohol related causes increased over 60% in both sexes between the late 1950s and the late 1970s. In absolute terms, this corresponds to about 10 000 more deaths per year in the later 1970s and is thus second only to the increase in tobacco related causes of death over the same calendar period. Alcohol related diseases represented a much larger

Fig 2 Standardised mortality ratios from liver cirrhosis in various Italian provinces, 1975-77: (a) males; (b) females.

proportion of all causes of deaths in Italy than in most other developed countries (with the sole exception of France)^{5,12} and, in particular, alcohol related neoplasms accounted for almost 9% of all cancer deaths.

There are, of course, several problems and uncertainties in the interpretation of the data presented. We tried to reduce bias related to the reliability of death certification by restricting our inferences on age standardised rates below age 65 and age specific ones below age 75, and independent studies and analyses of comparative completeness have indicated that Italian death certification in these age groups is reasonably accurate for most causes of death.¹³ This may, however, not be the case for primary liver cancer, where misclassification may be common even in middle age.⁵

A major problem is posed by the fact that the diseases considered are caused by factors other than alcohol, chiefly tobacco for cancers of the mouth, pharynx, oesophagus or larynx and hepatitis B virus for cancer or cirrhosis of the liver.^{5,14-16} Since there is a positive interaction between alcohol and tobacco on the excess risk of neoplasms of the upper digestive and respiratory tract,¹⁶ it is difficult to estimate precisely

the proportion of alcohol related excess deaths in the absence of tobacco. This interaction probably accounts also for the large male/female differences in mortality from these neoplasms, since smoking (and, in particular, pipe or cigar smoking) was extremely uncommon in Italian females over past decades.¹⁷ However, alcohol alone has been proved carcinogenic on human oesophagus.¹⁸ Further, it is known that modification in exposure to only one of these two risk factors can markedly modify the excess risk for subjects exposed to both.⁵ Similar considerations may be applied to the interaction between alcohol and hepatitis B virus in cirrhosis or carcinoma of the liver.^{14,19}

However complex the problems of interpretation, the data still indicate that mortality from all major alcohol related causes (whatever the influence of other aetiological factors) has largely increased in Italy during a period when average per capita alcohol consumption has roughly trebled. This finding is in general agreement with previous studies of trends in alcohol consumption and in mortality from various alcohol related causes in Britain or in Finland.²⁰⁻²²

Further evidence of an important role of alcohol in the mortality trends discussed is given by the

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inspection of age specific rates. In fact, marked upward trends were observed in mortality from all the alcohol related causes in young males, but death rates from cancers of the mouth or pharynx, oesophagus and larynx were declining for males born between 1900 and 1915, who were young adults in a period (between 1920 and 1935) when alcohol consumption was reduced in Italy.² This indirectly indicates an early stage effect of alcohol on carcinogenesis of the upper digestive and respiratory tract.²³

The geographical differences in mortality rates observed in various Italian provinces give additional evidence of a determinant role of alcohol, since mortality was grossly elevated in areas with higher alcohol consumption,¹⁰ and death certification rates from the major alcohol related causes were positively and (chiefly in males) strongly correlated. In some areas of north eastern Italy, deaths from liver cirrhosis alone accounted for over 18% of total years of work lost in males and about 10% of females (although the prevalence of serum hepatitis B markers in the same areas was lower than the national average),^{24,25} and mortality rates from cancer of the oesophagus in males were over five times the national average¹¹ and among the highest reported in western Europe.¹² In summary, the evidence presented indicates that the health consequences of heavy alcohol drinking in Italy are extremely severe and have been markedly increasing over the last three decades, in line with a large increase in alcohol consumption.

This work was conducted within the framework of the CNR (Italian National Research Council) Applied Projects "Oncology" (Contract No. 85.02209.44) and "Preventive and Rehabilitative Medicine" (Contracts No. 84.02233.56 and No. 84.02299.56).

The contributions of the Italian Association for Cancer Research and of the Italian League against Tumours, Milan, Italy, are gratefully acknowledged.

We thank Dr Alessandro Barchielli for providing figures on alcohol consumption, Dr Eva Negri for skilful help with data processing, and the Gustavus A Pfeiffer Memorial Library and Ms Antonietta Di Bitetto for editorial assistance.

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