



Original article

Hospital admissions and mortality in the 15 years after a first-time hospital contact with an alcohol problem: a prospective cohort study using the entire Danish population

Gro Askgaard (6), 1,2* David A Leon, 3,4 Thomas Deleuran and Janne S Tolstrup

¹National Institute of Public Health, University of Southern Denmark, Copenhagen K, Denmark, ²Gastro Unit, Copenhagen University Hospital, Bispebjerg Hospital, Copenhagen N, Denmark, ³Department of Non-Communicable Disease Epidemiology, Faculty of Epidemiology and Population Sciences, London School of Hygiene & Tropical Medicine, London, UK, ⁴Department of Community Medicine, UiT Arctic University of Norway, Tromsø, Norway and ⁵Department of Hepatology and Gastroenterology, Aarhus University Hospital, Aarhus C, Denmark

*Corresponding author. National Institute of Public Health, University of Southern Denmark, DK-1353 Copenhagen K, Denmark. E-mail: gask@dadlnet.dk

Editorial decision 21 June 2019; Accepted 11 July 2019

Abstract

Background: Potential benefits of preventing continued alcohol intake in individuals presenting at the hospital with an alcohol problem can be highlighted by studying their excess risk of subsequent morbidity and mortality.

Methods: All Danish residents with a first-time hospital contact with alcohol problems (intoxication, harmful use or dependence) in 1998–2002 were followed through 2012 using healthcare registries. We compared their cause-specific rates of hospital admission and mortality to the expected rates derived from the general population by calculating standardized incidence rate ratios.

Results: The 26716 men and 12169 women who were hospitalized with alcohol problems (median age 44 years) had more than 10 times the rate of subsequent admission to psychiatric departments and three times the rate of subsequent admission to somatic departments compared with the general population. In particular, the hospital admission rates for gastroenterological disease and injuries were high. The cumulative all-cause 10-year mortality risk was 29% [95% confidence interval (CI), 28–30] in men and 26% (95% CI, 24–27) in women with alcohol problems. The ratios of observed to expected death rate for all-cause mortality were 4.0 (95% CI, 3.8–4.1) in men and 4.3 (95% CI, 4.0–4.7) in women and, for causes of death fully attributable to alcohol, 16 (95% CI, 15–17) in men and 33 (95% CI, 29–38) in women.

Conclusions: Individuals hospitalized with alcohol problems have much higher rates of subsequent alcohol-related hospital admission and mortality than the general

population. Increased focus on preventing continued alcohol consumption in these individuals may reduce their subsequent morbidity and mortality.

Key words: Alcohol problems, public health, hospital admission, mortality, cohort study

Key Messages

- Men and women with a first-time hospital contact with obvious alcohol problems had a much higher rate of subsequent hospital admission and mortality due to a range of medical causes compared with the general population when followed up for more than 10 years.
- Much of this excess in subsequent hospital admission and mortality was from alcohol-related causes (mental and gastroenterological disease, injuries, fractures and poisonings).
- Overall, nearly a third of all deaths in men and one-quarter of all deaths in women were fully attributed to alcohol (caused by alcoholic liver disease, alcohol poisoning, etc.) in individuals with a hospital contact with alcohol problems.
- These findings show the ongoing alcohol problems among individuals hospitalized with alcohol problems in the years after the initial hospital contact with alcohol problems. Interventions to reduce alcohol consumption at initial hospital contact should be prioritized.

Introduction

Alcohol use is estimated to result in more than 2 million deaths and 85 million disability-adjusted life-years lost globally each year. Alcohol problems and dependency are often untreated, though reduction in alcohol consumption among problem drinkers decreases morbidity, mortality and healthcare costs. 5,6

Around 2% of all Danish hospital contacts involve alcohol diagnoses, reflecting obvious alcohol problems. ⁷⁻⁹ Despite the fact that the hospital setting is acknowledged as a unique opportunity for brief interventions and initiation of alcohol treatment, ¹⁰⁻¹² healthcare providers do not exploit this opportunity. ¹³⁻¹⁷ A US study revealed that only half of hospital patients with recognized alcohol problems received alcohol intervention or referral for treatment. ¹⁸ In the UK, where some hospitals even have alcohol care teams, only 60% of patients with recognized alcoholic liver disease were given advice or support for alcohol problems. ¹⁹

The potential benefit of actively targeting intervention and treatment towards patients who present with alcohol problems at the hospital is best understood by quantifying their subsequent risk of morbidity and mortality. If such patients are at especially high risk, the argument speaks in favour of a much more directed, as an alternative to the prevalent more passive, approach.

Therefore, we conducted a nationwide study of all men and women in Denmark who had a first-time hospital contact with alcohol problems between 1998 and 2002. We followed them for up to 15 years and estimated the incidence rates of cause-specific hospital admission and mortality compared with the general Danish population.

Material and methods

This was a nationwide, registry-based, prospective cohort study of all individuals with an initial hospital contact with alcohol problems between 1998 and 2002 in Denmark. The cohort was followed for subsequent hospital admissions and mortality until 2012.

Individuals with a hospital contact with alcohol problems

We included all individuals aged 20-84 years who went to the hospital for the first time with obvious alcohol problems during 1998-2002 in Denmark [alcohol intoxication (ICD-10: F10.0), harmful alcohol use (ICD-10: F10.1) or alcohol dependence (ICD-10: F10.2, F10.3, F10.4 and F10.5)]. 8,9 We identified hospital contacts with alcohol problems from all hospital departments (psychiatric, somatic, admission, emergency room and outpatient clinics). The diagnosis of alcohol problems could be the primary diagnosis or any other diagnosis listed for the hospital contact. Patients were excluded if they had a recorded hospital contact with alcohol problems prior to 1998 (identified from psychiatric hospital records from 1970 to 1998 and somatic hospital records from 1977 to 1998). We also excluded patients over 85 years of age because we lacked comparative data in 5-year age groups for individuals over 85 years.

All 5.6 million Danish citizens have access to universal, tax-financed healthcare.²⁰ Data on hospital contacts and diagnoses were obtained from the National Patient Register and the Danish Psychiatric Central Research Register.

The National Patient Register was established in 1977 and contains data on all somatic admissions, with emergency and outpatient contacts added in 1995. The Danish Psychiatric Central Research Register was established in 1970 and contains data on all psychiatric hospital admissions. Emergency and outpatient contacts were added to both hospital registers in 1995. Information on causes of death was obtained from the Danish Register of Causes of Death established in 1970. In Danish healthcare registries, diagnoses are recorded according to the eighth and, since 1994, the tenth International Classification of Diseases (ICD).

The level of comorbidity was computed from hospital diagnoses given prior to or at the time of the first-time hospital contact with alcohol problems. Somatic comorbidity was estimated according to the Charlson comorbidity index score.²⁴ Psychiatric comorbidity was estimated as the number of the major psychiatric disease categories: dementia and organic disorders not caused by alcohol (ICD-8: 290, 292, 294; ICD-10: F00-09), schizophrenia (ICD-8: 295, 298; ICD-10: F20-29), mood disorders (ICD-8: 296; ICD-10: F30-39), neurotic and stress-related disorders (ICD-8: 300; ICD-10: F40-49), behavioural syndromes associated with physiological disturbances (ICD-10: F50-59), personality disorders (ICD-8: 301; ICD-10: F60-69), mental retardation (ICD-10: F70-79), psychological development disorders (ICD-10: F80-89) and behavioural and emotional disorders (ICD-10: F90-99).

Information on marital status, vital status and migration was obtained from the Danish Civil Registration System²⁰ and information on education level came from Statistics Denmark. All the above registries were linked by a personal identification number: a unique identifier assigned to all Danish residents since 1968.

General population data for comparison

We used publicly available data aggregated by sex, age group and calendar year for cause-specific hospital admission and mortality for the general Danish population. Mortality rates by cause, sex, age group and calendar year for the Danish population were derived from the mortality database held by the World Health Organization. ²⁷

Follow-up for subsequent hospital admissions and cause of death

Individuals with a hospital contact with alcohol problems were followed for all subsequent hospital admissions (inpatient admissions) and death. The cohort was followed from the date of termination of first-time hospital contact with alcohol problems until death (n = 13088), reaching age 85 years (n = 628) or end of follow-up (31 December 2012),

whichever occurred first. To ensure comparability with data derived from the general Danish population, we used the primary diagnosis listed as the cause of hospital admission and the underlying cause for cause of death. ICD-10 codes for cause-specific hospital admissions and causes of death used in this paper are listed in Supplementary Appendix A, available as Supplementary data at *IJE* online.

Statistical analyses

Indirect standardization was used to compare observed numbers of subsequent hospital admissions and deaths with those expected based on age, sex and calendar-year-specific rates in the total general population. The ratio of observed to expected events (O/E) provides an estimate of the relative risk for hospital admissions and mortality in individuals with a hospital contact with alcohol problems compared with the general population. By definition, the general population used includes those included in our cohort. The observation time was from the date of termination of the first-time hospital contact with an alcohol problem to death, reaching age 85 years or end of follow-up, whichever was the soonest. Days spent in hospital did not contribute to the observation time. Poisson regression was used to model standardized incidence ratios (SIRs) or standardized mortality ratios (SMRs) for hospital admissions and mortality in men and women with a hospital contact with alcohol problems relative to the general population.

People who drink heavily also tend to smoke, which increases their morbidity and mortality. ^{1,28} Based on a prior suggested definition, we calculated causes of death mostly attributed to tobacco. ²⁹ We also measured the 'excess' of subsequent hospital admissions related to alcohol in individuals hospitalized with alcohol problems compared with the general population. These non-smoking causes of hospital admission were mental, gastroenterological and external causes of disease (injuries, fractures and poisonings). ⁹ We calculated the proportion that these non-smoking causes (NS causes) contributed to the excess of the total number (TN) of hospital admissions to somatic departments in men and women with a hospital contact with alcohol problems compared with the general population:

[Observed (NS) – Expected (NS)]/ [Observed (TN) – Expected (TN)] * 100%

Analyses were carried out using Stata (version 15; Stata-Corp LP, College Station, TX).

Ethics

All data were delivered anonymized. The Danish Data Protection Agency approved the study (j.nr. 2014–41-3516).

Results

Between 1998 and 2002, 26716 men and 12169 women in Denmark had a first-time hospital contact where they were given an alcohol-problem diagnosis (Table 1). Alcohol-problem diagnoses for intoxication and dependence were equally common (38–42%), whereas 21–22% had a diagnosis for harmful alcohol use. The majority had their hospital contact at a somatic department. Median age (interquartile range; IQR) was 44 years (33–55) in men and 45 years (34–56) in women. About half the patients had 9 or fewer years of education and the majority were unmarried. At the time of the hospital contact, the majority of individuals had no somatic or psychiatric comorbidity. Seven out of 10 had a Charlson comorbidity index score of zero and the vast majority had not been registered with a psychiatric diagnosis.

Overall rate of subsequent hospital admissions

During the follow-up, 391 003 person-years (mean follow-up 10 years) were accumulated, during which we observed 50 179 hospital admissions to psychiatric departments and 226 885 to somatic departments among individuals with a first-time hospital contact with alcohol problems (Table 2). Compared with the general population, men and women hospitalized with alcohol problems had a more than 10 times higher rate of subsequent hospital admissions in psychiatric departments {SIR in men, 12 [95% confidence interval (CI), 11–12], SIR in women, 16 [95% CI, 16–17]} and about a three times higher rate of hospital admission in somatic departments [SIR in men, 3.0 (95% CI, 3.0–3.0), SIR in women, 2.8 (95% CI, 2.8–2.9)].

The rate of outpatient visits to somatic departments (calculated for 2006–2013) was slightly higher for men with a hospital contact with alcohol problems compared with men in the general population [SIR, 1.2 (95% CI, 1.2–1.2)], but this difference was not found among women [SIR, 1.0 (95% CI, 1.0–1.0)].

Cause-specific subsequent hospital admission

Men and women with a hospital contact with alcohol problems had higher rates of subsequent hospital admissions for all diagnoses compared with the general population (Table 2). The diagnosis-specific SIRs were mostly similar in men and women, though they were higher in women than men for the alcohol-related mental and behavioural disorders, liver and pancreatic disease and for poisoning by alcohol or other substances. In both sexes, it was among these alcohol-related diseases that the highest SIRs were found. For example, the SIR for hospital

Table 1. Characteristics in men and women with a hospital contact with alcohol problems between 1998 and 2002 ($n = 38\,885$). Values are numbers (percentages) unless stated otherwise

	Men	Women
	26 716 (69)	12 169 (31)
Characteristics of the initial	hospital contact with	alcohol problems
Alcohol diagnosis		
Alcohol intoxication	10 300 (39)	4835 (40)
Harmful alcohol use	5485 (20)	2692 (22)
Alcohol dependence	10 931 (41)	4642 (38)
Department		
Somatic	20 680 (77)	8760 (72)
Psychiatric	6036 (23)	3409 (28)
Type of hospital care		
Admission	14 144 (53)	6280 (52)
Emergency	8477 (32)	4013 (33)
Outpatient	4095 (15)	1876 (15)
Demographic and medical of	characteristics	
Age, median (IQR)	44 (33-55)	45 (34-56)
Education, years		
≤9	13 007 (49)	6753 (55)
10–11	11 181 (42)	3234 (27)
≥12	2528 (9.4)	2182 (18)
Civil status		
Never married	12 840 (48)	4225 (35)
Married	7833 (29)	4217 (35)
Other	5478 (21)	3497 (29)
Missing	565 (2.1)	230 (1.8)
Charlson comorbidity index	x, score	
0	18 636 (70)	8501 (70)
1	3927 (15)	1805 (15)
≥2	4153 (15)	1803 (15)
Psychiatric comorbidity,		
number of diagnoses		
0	23 044 (86)	8911 (73)
1	2198 (8.2)	1772 (15)
≥2	1474 (5.5)	1486 (12)

admission due to liver and pancreatic disease was 11 (95% CI, 11–12) in men and 17 (95% CI, 15–18) in women with alcohol problems compared with the general population.

SIRs ranged from three to five for admission for stroke, respiratory disease, nervous-system disease, gastro-enterological disease, infectious disease, ill-specified disease, admission for observation and for admission with fractures and injuries. SIRs ranged from one and a half to two for hospitalization due to cancer, cardiovascular disease, ischemic heart disease and musculoskeletal disease.

NS causes of hospital admission (mental, gastroenterological and external causes of disease, i.e. injuries, fractures and poisonings) accounted for 46% (men) and 45% (women) of all the 'excess' observed admissions in the

Table 2. Observed numbers (O), expected numbers (E) and standardized incidence ratios (SIRs) of subsequent hospital admissions according to diagnosis in men and women with a hospital contact with alcohol problems in the years between 1998 and 2002 (n = 38 885)

Diagnosis		Men		Women			
	О	Е	SIR (95%)	О	Е	SIR (95%)	
Hospital admission to psychiatric departments							
Any diagnosis ¹	30 699	2488	12 (11-12)	19 480	1087	16 (16-17)	
Mental disease due to alcohol or substance abuse ¹	10 614	563	15 (15-16)	5225	126	46 (43-48)	
Outpatient visit to somatic departments							
Any diagnosis ²	182 651	152 904	1.2 (1.2-1.2)	104 027	100 463	1.0 (1.0-1.0)	
Hospital admission to somatic departments							
Any diagnosis	147 360	48 604	3.0 (3.0-3.0)	79 525	27 881	2.8 (2.8-2.9)	
Mental and behavioral disease	17 764	1130	15 (15-16)	7720	294	26 (25-28)	
Cancer	6356	4240	1.5 (1.4-1.5)	3600	2375	1.7 (1.6-1.8)	
Cardiovascular disease	12 378	7207	1.7 (1.7-1.8)	3623	1902	2.0 (1.8-2.1)	
Ischemic heart disease	4185	3055	1.4 (1.3-1.5)	923	639	1.7 (1.5-2.0)	
Stroke	2813	1075	2.6 (2.4-2.8)	1264	388	3.0 (2.6-3.3)	
Respiratory disease	6700	2107	3.3 (3.2-3.5)	3824	1018	3.5 (3.2-3.7)	
Nervous system disease	4010	1423	2.8 (2.7-3.0)	1896	627	3.2 (2.9-3.5)	
Gastroenterological disease	17 576	4209	4.1 (4.0-4.3)	7865	1631	5.0 (4.8-5.3)	
Liver and pancreatic disease	7820	705	11 (11-12)	3242	207	17 (15-18)	
Infectious disease	9827	3505	2.7 (2.6-2.8)	4606	1469	3.0 (2.8-3.2)	
Musculoskeletal disease	5574	2928	1.7 (1.6-1.8)	3009	1614	1.7 (1.6-1.9)	
Ill specified disease	11 323	3718	2.8 (2.7-3.0)	5942	1936	3.0 (2.8-3.2)	
Admission for observation	11 157	3892	2.7 (2.5-2.8)	5743	1891	3.1 (2.9-3.3)	
External causes	20 184	5073	3.8 (3.7-4.0)	11917	2085	5.4 (5.2-5.7)	
Fractures	5522	1506	3.7 (3.5-3.9)	3207	753	4.3 (4.0-4.6)	
Injuries	10 484	3092	3.3 (3.2-3.4)	4218	1058	4.0 (3.8-4.3)	
Poisoning by alcohol or other substances	4169	476	7.7 (7.2-8.3)	4476	273	15 (14-16)	

SIRs are adjusted for calendar year of hospital contact with alcohol problems.

alcohol problem cohort compared with the general population (which also included those hospitalized with alcohol problems).

All-cause mortality

During follow-up of the cohort, 9451 men and 3637 women died. The cumulative all-cause mortality in men and women was 5.2% (95% CI, 4.0–5.8) and 3.8% (95% CI, 3.0–4.5) after 1 year, 16% (95% CI, 15–17) and 14% (95% CI, 13–15) after 5 years and 29% (95% CI, 28–30) and 26% (95% CI, 24–27) after 10 years (Figure 1). The SMR for all-cause mortality was 4.0 (95% CI, 3.8–4.1) in men and 4.3 (95% CI, 4.0–4.7) in women with a hospital contact with alcohol problems compared with the general population (Table 3). SMRs were highest in those who were young and middle-aged (20–49 years) at the time of the hospital contact with alcohol problems. For example, in men with alcohol problems, the SMR was 7.2 (95% CI, 6.7–7.7) for 30–39 years and

2.1 (95% CI, 1.9–2.2) for 70–84 years compared with the general population.

Cause-specific mortality

Overall, 2703 (29%) of all deaths in men and 915 (25%) of all deaths in women in the cohort were fully attributed to alcohol (caused by alcoholic liver disease, alcohol poisoning, etc.) (Table 4). Comparing rates of causes of deaths to the general population, SMRs for the cohort were higher for nearly all causes of death investigated for men and women.

SMRs for death due to mental disease caused by alcohol or substance abuse, liver disease and the aggregate of all causes fully attributable to alcohol were higher in women than men. For example, the SMR for all causes fully attributable to alcohol was 16 (95% CI, 15–17) in men but 33 (95% CI, 29–38) in women. For all other causes, the SMRs were relatively similar between the sexes. In both men and women, it was among these alcohol-related

^aPsychiatric admissions were calculated for 2000-2013 due to limited data availability.

^bSomatic outpatients visits were calculated for 2006–2013 due to limited data availability.

causes that the highest SMRs were found (ranging from 10 to 34).

SMRs were three to six times greater in the majority of causes of death examined, including the aggregate of all causes mostly attributable to tobacco, cancers of the upper aerodigestive tract (mouth, throat, upper oesophagus, etc.), liver cancer, cardiovascular disease, cardiomyopathy, stroke, respiratory disease, infectious disease, musculoskeletal disease in women, ill-specified diseases, external causes, suicide and accidents in men (see Supplementary Appendix A, available as Supplementary data at *IJE* online, for definitions). For example, the aggregate of causes mostly attributable to tobacco showed SMRs of 3.5 (95% CI, 3.1–4.0) in men and 3.5 (95% CI, 2.9–4.2) in women.

Discussion

This study defined the cohort of all Danish men and women who had a first-time hospital contact with alcohol problems between 1998 and 2002 and has examined their subsequent rate of hospital admissions and mortality

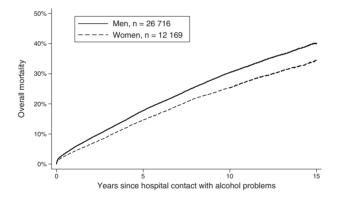


Figure 1. Cumulative all-cause mortality for men and women according to years since hospital contact with alcohol problems, in the years between 1998 and 2002 (n = 38.885).

through to 2012. We found that such patients had much higher rates of hospital admission than the Danish population as a whole: 10 times higher to psychiatric departments and three times higher to somatic departments. Cumulative mortality risks approximated one-third after 10 years. Hospital admission and mortality rates were higher for a range of medical conditions, in particular those associated with heavy drinking, including mental, gastroenterological and external causes of disease.

Their high mortality and morbidity in the subsequent 15 years after a hospital contact with alcohol problems indicate ongoing health problems most likely due to alcohol among these patients.

Strengths of our study include its nationwide setting with complete data on hospital care and causes of death. ^{20,21,23} The validity of hospital-discharge diagnoses is high: the positive predictive value of the primary hospital-discharge diagnosis in the registry was 81% when compared with a medical chart review of a random sample of 1000 admissions. ²¹ Thus, our findings are likely to represent valid populations-based estimates on the morbidity and mortality of patients hospitalized with alcohol problems in Denmark.

Limitations of our study include the inability to separate the effects of heavy drinking from other factors such as smoking and socio-economic position on the risk of admissions and mortality. However, NS causes (mental, gastroenterological and external) of hospital admissions accounted for almost half of all the 'excess' hospital admissions observed compared with the general population. For mortality, SMRs were much greater for causes fully attributable to alcohol than causes mostly attributable to tobacco. Therefore, although smoking may explain some of the excess morbidity and mortality, it is unlikely to alter our conclusion that ongoing alcohol use in these individuals with a hospital contact with alcohol problems causes a high burden of hospital admissions and mortality.

Table 3. Observed numbers (O), expected numbers (E) and standardized mortality ratios (SMRs) in men and women according to age at hospital contact with alcohol problems, in the years between 1998 and 2002 ($n = 38\,885$)

Age at hospital contact with alcohol problems	Men				Women			
	Person-years	О	Е	SMR (95% CI)	Person-years	О	E	SMR (95% CI)
Overall, 20–84	265 451	9200	2206	4.0 (3.8–4.1)	125 551	3452	<u>757</u>	4.3 (4.0–4.7)
20-29	66 106	345	57	5.5 (5.0-6.2)	29 218	66	11	5.6 (4.4-7.2)
30-39	62 989	991	126	7.2 (6.7–7.7)	24 822	286	30	8.8 (7.8-10)
40-49	61 188	2051	320	5.9 (5.5-6.2)	32 824	750	109	6.4 (5.8–7.1)
50-59	46 330	2553	546	4.3 (4.0-4.5)	23 800	954	185	4.8 (4.4-5.2)
60-69	20 747	1962	595	3.0 (2.8-3.2)	10 138	785	198	3.7 (3.3-4.0)
70-84	8091	1298	562	2.1 (1.9–2.2)	4750	611	225	2.5 (2.3–2.8)

Table 4. Observed numbers (O), expected numbers (E) and standardized cause-specific mortality ratios (SMRs) in men and women with a hospital contact with alcohol problems, in the years between 1998 and 2002 (n = 38885)

	Men			Women		
Cause of death	О	E	SMR (95% CI)	О	E	SMR (95% CI)
Mental disorders	1294	12	13 (11–14)	416	24	18 (15–22)
Mental disease caused by alcohol or substance abuse	1162	72	15 (14–17)	331	10	34 (28-43)
Cancer	1622	731	2.1 (1.8-2.3)	722	314	2.0 (1.7-2.3)
Upper aerodigestive tract cancer (mouth, throat, upper oesophagus, etc.)	64	350	5.1 (4.1-6.5)	10	92	6.4 (3.9–11)
Lung cancer	454	190	2.2 (1.8-2.8)	231	77	2.5 (1.9-3.3)
Stomach, colon and rectal cancer	193	113	1.4 (1.0-1.9)	61	41	1.5 (0.9–2.5)
Liver cancer	113	20	6.3 (4.3-9.2)	26	4	4.9 (2.1-12)
Pancreatic cancer	68	41	1.1 (0.6-2.1)	34	17	1.5 (0.7–3.3)
Breast cancer	<5	< 5	N/A	92	56	1.4 (0.9-2.2)
Cardiovascular disease	1618	57	3.1 (2.8-3.4)	497	143	3.0 (2.5-3.7)
Acute myocardial infarction	36	133	2.0 (1.6-2.6)	88	30	2.7 (1.7-4.2)
Ischemic heart disease other than Acute myocardial infarction	345	119	3.4 (2.8-4.2)	80	25	3.0 (1.9-4.7)
Cardiomyopathy	67	11	6.1 (3.7-10)	8	2	6.0 (1.5-24)
Stroke	470	129	3.2 (2.6-3.9)	178	50	2.9 (2.1-4.1)
Respiratory disease	737	130	5.7 (4.9-6.6)	366	62	5.7 (4.6–7.1)
Chronic obstructive pulmonary disease and asthma	484	113	4.7 (3.9-5.7)	283	57	4.4 (3.4–5.7)
Nervous system disease	152	54	2.6 (1.8-3.8)	63	23	2.8 (1.6-4.7)
Gastroenterological disease	1806	135	12 (11-13)	729	41	16 (14–19)
Pancreatitis	123	10	9.9 (6.5-15)	29	2	10 (4.4–25)
Liver disease	1430	80	16 (14–18)	574	19	29 (25-35)
Infectious disease	353	66	4.7 (3.7-6.0)	124	23	7.2 (5.2–10)
Musculoskeletal disease	19	8	2.3 (0.9-6.1)	18	6	3.0 (1.1–7.9)
Ill-specified disease	396	15	3.4 (2.7-4.2)	166	32	6.8 (5.1–9)
External causes of death	629	10	6.2 (5.3–7.3)	20	18	10 (7.6–14)
Suicide	381	58	6.4 (5.2–7.9)	138	10	12 (8.2–17)
Accidents	93	28	3.5 (2.3-5.3)	10	4	2.3 (1.2-4.3)
All causes fully attributable to alcohol ³⁰	2703	156	16 (15–17)	915	27	33 (29–38)
All causes mostly (≥74%) attributable to tobacco ²⁹	1268	363	3.5 (3.1-4.0)	592	142	3.5 (2.9–4.2)

SMRs are adjusted for calendar year of hospital contact with alcohol problems.

To the best of our knowledge, no prior studies have assessed the frequency of hospital admissions due to a range of causes in individuals hospitalized with alcohol problems compared with the general population. Two prior studies of patients hospitalized with alcohol problems found higher incidences of later readmission for liver cirrhosis, pancreatitis, cancer and accidents compared with controls. 31,32

Our reported SMRs for all-cause and cause-specific mortality for men and women hospitalized with alcohol problems compared with the general population are in line with previous meta-analyses of alcohol-treatment cohorts. 5,33 Half the patients in our study had a diagnosis of less severe alcohol problems (alcohol intoxication and harmful alcohol use) and only 40% were diagnosed with alcohol dependence. Nevertheless, we found a high rate of alcohol-related hospital admission and deaths. These observations are consistent with a study that followed more than 2000 individuals after an index alcohol-related healthcare contact and found a high number of individuals with recurrent

alcohol-related contacts even among those individuals diagnosed with alcohol intoxication at their index alcohol contact (20% had at least one contact per year in the following 10 years) and harmful alcohol use (40% had at least one contact per year in the following 10 years).⁸

More than 25% of the observed deaths in our study were directly attributable to alcohol and this proportion is most likely conservative. ^{34,35} In a validation study of 403 deaths, the proportion of deaths where the cause was attributable to alcohol rose from 25% to 36% when information from medical records and autopsies was added to registry data. ³⁵ Further, the high SMRs of death due to unspecified causes and ischemic heart disease (other than myocardial infarction) may reflect mortality due to alcohol, which was observed to fluctuate with alcohol-induced death in Russia. ³⁶

Those most at risk of subsequent hospital admission are also likely to be those most at risk of dying. To this extent, hospital-admission rates in the cohort might have been even higher if mortality had been postponed. However, as the focus of this paper considers both outcomes as indicators of disease burden in the cohort, our analyses in this respect are not biased, being based on complete follow-up of the whole population.

Heavy drinking is associated with psychiatric disease ^{37,38} and increases the risk of more than 200 diseases including cancers, heart disease, neurologic disease, infectious disease and gastrointestinal diseases, in particular cirrhosis and pancreatitis. ³⁹ With respect to cancer, the risk was five-fold increased for mouth, throat and oesophagus cancer, but only slightly increased for colon cancer and breast cancer in women. This observation is in accordance with the fact that alcohol explains about 40% of mouth, throat and oesophagus cancer cases (the attributable fraction), but only 10–15% of colon cancer cases and 7% of breast cancer cases. ⁴⁰

Around 620 000 individuals (14%) in Denmark have harmful alcohol use and 150 000 individuals (3%) are alcohol-dependent. With only 20 000 individuals receiving specialized alcohol treatment, there is substantial undertreatment of alcohol problems in Denmark, in line with the situation in other countries.^{4,41}

This study points to a potential substantial reduction in healthcare costs, morbidity and premature mortality if interventions to reduce harmful alcohol consumption were prioritized among patients hospitalized for alcohol problems. The challenge to be addressed, however, is to develop optimal approaches to identify which hospitalized patients would benefit most from interventions to reduce their alcohol intake. Hospital contacts with alcohol problems represent an opportunity to raise these patients' awareness of the threat that alcohol intake poses to their health. 42–44 Observational data suggest that brief interventions and the implementation of alcohol care teams in the hospital setting reduce 1-year mortality, alcohol-related hospital admissions and healthcare cost, 30,45 although, ideally, such interventions should be studied in randomized–controlled trials.

In conclusion, men and women hospitalized with alcohol problems have much higher rates of subsequent alcohol-related hospital admission and mortality than the general population. Thus, disease and premature mortality may be reduced by increased focus on preventing continued alcohol consumption in these individuals.

Supplementary Data

Supplementary data are available at IJE online.

Funding

This study received no funding.

Acknowledgements

All authors designed the study. G.A. performed the analyses and drafted the manuscript. All authors contributed to discussing and revising the paper.

Conflict of interest: All authors have completed the ICMJE uniform disclosure form at www.icmje.org/coi_disclosure.pdf and declare: no support from any organization for the submitted work; no financial relationships with any organizations might have an interest in the submitted work in the previous 3 years; no other relationships or activities that could appear to have influenced the submitted work.

References

- Forouzanfar MH, Afshin A, Alexander LT et al. Global, regional, and national comparative risk assessment of 79 behavioural, environmental and occupational, and metabolic risks or clusters of risks, 1990–2015: a systematic analysis for the Global Burden of Disease Study 2015. Lancet 2016;388: 1659–724.
- Alonso J, Angermeyer MC, Bernert S et al. Use of mental health services in Europe: results from the European Study of the Epidemiology of Mental Disorders (ESEMeD) project. Acta Psychiatr Scand Suppl Scand 2004;109:47–54.
- 3. Kohn R, Saxena S, Levav I, Sraceno B. The treatment gap in mental health care. *Bull World Health Organ* 2004;82:858–66.
- 4. Rehm J, Manthey J, Struzzo P, Gual A, Wojnar M. Who receives treatment for alcohol use disorders in the European Union? A cross-sectional representative study in primary and specialized health care. *Eur Psychiatry* 2015;30:885–93.
- Roerecke M, Rehm J. Alcohol use disorders and mortality: a systematic review and meta-analysis. Addiction 2013;108: 1562–78.
- Charlet K. Harm reduction-a systematic review on effects of alcohol reduction on physical and mental symptoms. Addict Biol 2017;22:1119–59.
- 7. Danish Health and Medicines Authority, Statens Serum Institut. Alkoholstatistik 2015. Nationale data, 2015.
- 8. Ahacic K, Damström-Thakker K, Kåreholt I. Recurring alcoholrelated care between 1998 and 2007 among people treated for an alcohol-related disorder in 1997: a register study in Stockholm County. *BMC Public Health* 2011;11:574.
- The World Health Organization. International Classification of Diseases, 10th Edn (ICD-10). Geneva: The World Health Organization, 2010.
- National Institute for Health and Care Excellence. Alcohol-use disorders: prevention. NICE Clin Guidel, June 2010.
- World Health Organization. European Status Report on Alcohol and Health 2010. Copenhagen: World Health Organization, 2010.
- 12. OECD. Tackling Harmful Alcohol Use: Economic and Public Health Policy. Paris: OECD Publishing, 2015.
- McKnight-Eily RL, Liu Y, Brewer DR et al. Vital signs: communication between health professionals and their patients about alcohol use—44 states and the district of Columbia, 2011.
 MMWR Recomm Rep 2014;63:16–22.
- 14. Wood E, Samet JH, Volkow ND. Physician education in addiction medicine. *JAMA* 2013;310:1673–74.

- 15. Kelly JF, Wakeman SE, Saitz R. Stop talking 'dirty': clinicians, language, and quality of care for the leading cause of preventable death in the United States. *Am J Med* 2015;128:8–9.
- 16. Sagar N, Dibor C, Polson R. An audit to evaluate the use of the alcohol fast screening tool in acute medical admissions in a district hospital. *Gut* 2013;62:A12–13.
- 17. Mayor S. Hospitals miss chances to cut deaths from alcohol related liver disease, report says. *BMJ* 2013;346:f3781.
- Smothers BA, Yahr HT, Ruhl CE. Detection of alcohol use disorders in general hospital admissions in the United States. *Arch Intern Med* 2004;164:749–56.
- 19. Measuring the Units—a Review of Patients Who Died with Alcohol-related Liver Disease. London: National Confidential Enquiry into Patient Outcome and Death, 2013.
- Schmidt M, Pedersen L, Sørensen HT. The Danish Civil Registration System as a tool in epidemiology. Eur J Epidemiol 2014;29:541–49.
- Schmidt M, Schmidt SAJ, Sandegaard JL, Ehrenstein V, Pedersen L, Sørensen HT. The Danish National Patient Registry: a review of content, data quality, and research potential. *Clin Epidemiol* 2015;7:449–90.
- 22. Mors O, Perto GP, Mortensen PB. The Danish Psychiatric Central Research Register. *Scand J Public Health* 2011;39:54–57.
- 23. Helweg-Larsen K. The Danish Register of Causes of Death. *Scand J Public Health* 2011;**39**:26–29.
- 24. Thygesen SK, Christiansen CF, Christensen S, Lash TL, Sørensen HT. The predictive value of ICD-10 diagnostic coding used to assess Charlson comorbidity index conditions in the population-based Danish National Registry of Patients. BMC Med Res Methodol 2011:11:83.
- 25. Statistics Denmark. Statbank. p. https://www.dst.dk/en/Statistik (9 March 2018, date last accessed).
- Danish Psychiatric Central Research Register. http://www.kea. au.dk/da/forskning/det-psykiatriske-centralregister.html (9 March 2018, date last accessed).
- World Health Organization. WHO mortality database. http://apps.who.int/healthinfo/statistics/mortalit (9 March 2018, date last accessed).
- 28. Askgaard G, Grønbæk M, Kjær MS, Tjønneland A, Tolstrup JS. Alcohol drinking pattern and risk of alcoholic liver cirrhosis: a prospective cohort study. *J Hepatol* 2015;62:1061–67.
- 29. Oza S, Thun MJ, Henley SJ, Lopez AD, Ezzati M. How many deaths are attributable to smoking in the United States? Comparison of methods for estimating smoking-attributable mortality when smoking prevalence changes. *Prev Med (Baltim)* 2011;52:428–33.
- Quality and Productivity: Proven Case Study. Alcohol Care Teams: Reducing Acute Hospital Admissions and Improving Quality of Care. Bolton: The British Society of Gastroenterology, 2014.

- Piette JD, Barnett PG, Moos RH. First-time admissions with alcohol-related medical problems: a 10-year follow-up of a national sample of alcoholic patients. J Stud Alcohol 1998;59:89–96.
- 32. Schwarzinger M, Thiébaut SP, Baillot S, Mallet V, Rehm J. Alcohol use disorders and associated chronic disease—a national retrospective cohort study from France. *BMC Public Health* 2018;18:43.
- Roerecke M, Rehm J. Cause-specific mortality risk in alcohol use disorder treatment patients: a systematic review and meta-analysis. *Int J Epidemiol* 2014;43:906–19.
- 34. Blake JE, Compton KV, Schmidt W, Orrego H. Accuracy of death certificates in the diagnosis of alcoholic liver cirrhosis. *Alcoholism Clin Exp Res* 1988;12:168–72.
- Ågren G, Jakobsson SW. Validation of diagnoses on death certificates for male alcoholics in Stockholm. Forensic Sci Int 1987; 33:231–41.
- 36. Zaridze D, Maximovitch D, Lazarev A *et al.* Alcohol poisoning is a main determinant of recent mortality trends in Russia: evidence from a detailed analysis of mortality statistics and autopsies. *Int J Epidemiol* 2009;38:143–53.
- Hasin DS, Tsai WY, Endicott J, Mueller TI, Coryell W, Keller M. Five-year course of major depression: effects of comorbid alcoholism. *J Affect Disord* 1996;41:63–70.
- 38. Flensborg-Madsen T, Mortensen EL, Knop J, Becker U, Sher L, Grønbaek M. Comorbidity and temporal ordering of alcohol use disorders and other psychiatric disorders: results from a Danish register-based study. Compr Psychiatry 2009;50: 307–14.
- Rehm J, Gmel GE, Gmel G et al. The relationship between different dimensions of alcohol use and the burden of disease—an update. Addiction 2017:112:968–1001.
- Praud D, Rota M, Rehm J et al. Cancer incidence and mortality attributable to alcohol consumption. Int J Cancer 2016;138: 1380–87.
- 41. McLellan AT, Woodworth AM. The affordable care act and treatment for 'substance use disorders': implications of ending segregated behavioral healthcare. *J Subst Abuse Treat* 2014;46: 541–45.
- 42. Lid TG, Oppedal K, Pedersen B, Malterud K. Alcohol-related hospital admissions: missed opportunities for follow up? A focus group study about general practitioners' experiences. *Scand J Public Health* 2012;40:531–36.
- 43. Apodaca TR, Schermer CR. Readiness to change alcohol use after trauma. *J Trauma* 2003;54:990–94.
- 44. Sheron N, Moore M, O'Brien W, Harris S, Roderick P. Feasibility of detection and intervention for alcohol-related liver disease in the community: the Alcohol and Liver Disease Detection study (ALDDeS). Br J Gen Pract 2013;63:e698–705.
- Mcqueen J, Te H, Allan L, Mains D, Hardy V. Brief interventions for heavy alcohol users admitted to general hospital wards. *Cochrane Database Syst Rev* 2011;8:CD005191.