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# Social inequality and incidence of and survival from tumours of the central nervous system in a population-based study in Denmark, 1994–2003

Lisbeth Samsø Schmidt\*, Hanne Nielsen, Sven Schmiedel, Christoffer Johansen

Institute of Cancer Epidemiology, Danish Cancer Society, Strandboulevarden 49, DK-2100 Copenhagen Ø, Denmark

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## ABSTRACT

We investigated the effects of socioeconomic, demographic and health-related indicators on the incidence of and survival from tumours of the central nervous system (CNS) diagnosed in 1994–2003 with follow-up through 2006 in Denmark using information from nationwide Danish administrative registers. The analyses were based on data on 5622 patients with CNS tumours in a cohort of 2.7 million people born between 1925 and 1973 and aged  $\geq 30$  years. Socioeconomic and demographic factors were not associated with the incidence of CNS tumours, except for a significantly increased incidence rate ratio (IRR) amongst men in the agricultural class (IRR, 1.23; 95% CI, 1.04–1.45). The 1- and 5-year survival was significantly longer in higher socioeconomic groups, as assessed by education, income, affiliation to the work market and size of dwelling.

Socioeconomic position does not affect the incidence of CNS tumours but appears to be a prognostic factor for survival from CNS tumours in Denmark.

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## 1. Introduction

Tumours of the central nervous system (CNS) are rare; however, the incidence rates have increased during the past few decades in both men and women.<sup>1</sup> The increased incidence may be explained by improved diagnostic methods and by a lower threshold for performing diagnostic computed tomography or magnetic resonance image scanning; however, unknown risk factors may also play a role. In 2003, 922 new cases of CNS tumours were diagnosed in Denmark.<sup>2</sup> The results of studies of an association between incidence and socioeconomic position (SEP) are conflicting: several reported increased incidences of CNS tumours amongst persons with higher SEP,<sup>3–5</sup> whilst others found no association with SEP.<sup>6,7</sup>

Age and tumour characteristics are known to be important prognostic factors for survival,<sup>8</sup> but the effect of SEP has been

investigated only rarely.<sup>9–11</sup> The aim of this study was to investigate the effects of various socioeconomic indicators on the incidence of and survival from CNS tumours in Denmark on the basis of information from population-based, nationwide Danish administrative registers. It was carried out as part of a comprehensive, rigorous analysis of the role of socioeconomic position in cancer incidence and survival.

## 2. Material and methods

The material and methods are described elsewhere.<sup>12</sup> Briefly, the study population comprised all Danish residents born between 1925 and 1973 without a previous cancer and who entered the cohort at age 30 (see Fig. 1 in [12]). Information on socioeconomic, demographic and health-related indicators was obtained from various Danish administrative registers.<sup>12</sup> Crude, age-specific and age-standardised incidence rates

\* Corresponding author: Tel.: +45 35257500; fax: +45 35257731.

E-mail address: [samsoe@cancer.dk](mailto:samsoe@cancer.dk) (L.S. Schmidt).

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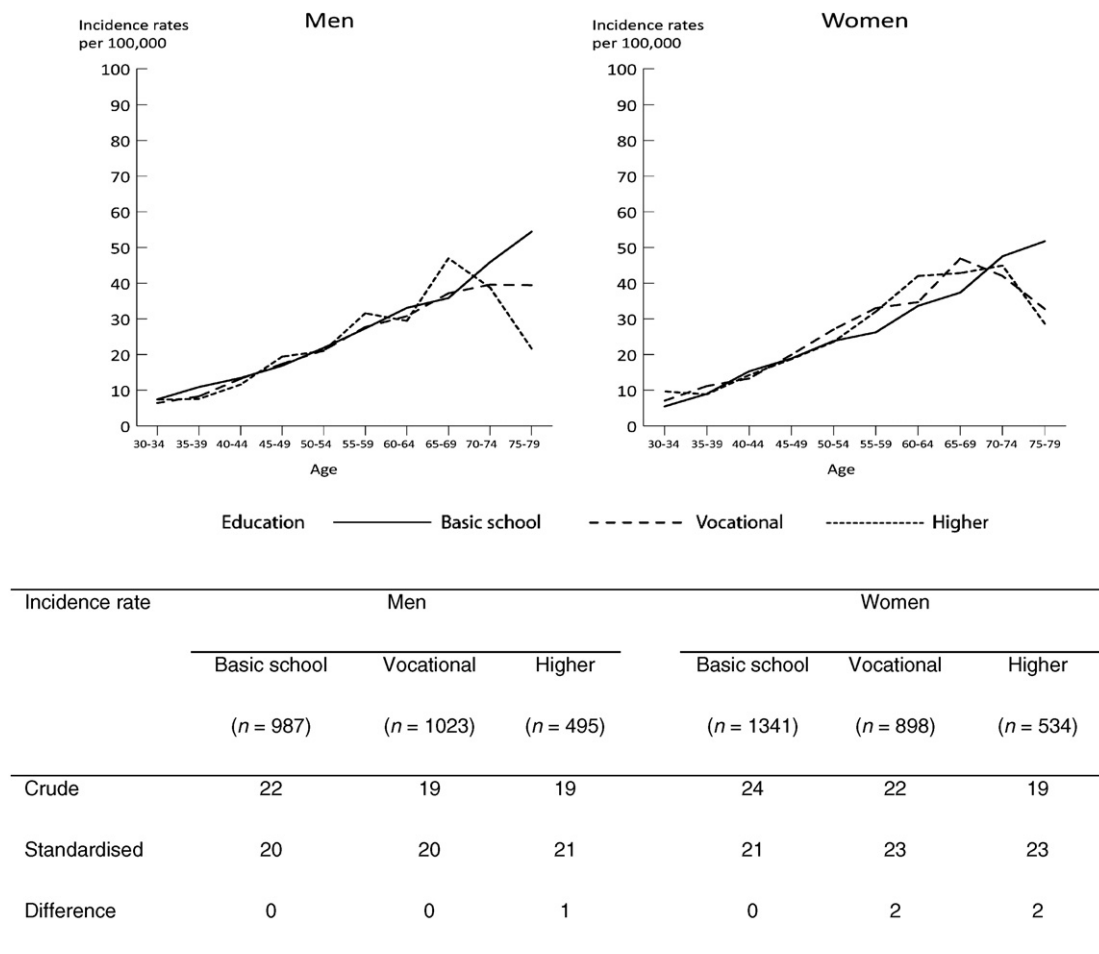
are presented for CNS tumours (ICD-10 C70–C72, D32–D33, D42–43) diagnosed in the cohort in 1994–2003. In a subset of the data, the tumours were restricted to malignant tumours of the CNS defined as described in the WHO classification.<sup>13</sup> The incidence rates were standardised by age (in 5-year age groups) and period (in two 5-year periods), with the total study population as the standard.<sup>14</sup> Further, we used log-linear Poisson regression to model incidence rate ratios (IRRs), first adjusted for period (in 5-year periods) and age (as two continuous variables: age and age<sup>2</sup> in years) and second by adding education and disposable income to the models. For each level of each indicator, we conducted relative survival analyses, adjusting for population mortality amongst the incident CNS tumour cases in 1994–2003 with follow-up through 2006.<sup>12</sup> Population mortality rates were stratified by age, period and the respective indicator. Except for the analyses of ethnicity, all analyses included only residents born in Denmark to at least one Danish-born parent with Danish citizenship.<sup>12</sup>

### 3. Results

We included 5622 persons in whom a CNS tumour was diagnosed during the study period 1994–2003, of which 2391 were malignant tumours. Amongst Danish persons, the male:female ratio for all CNS tumours was 0.90, and that for malignant tumours was 1.4. The age- and period-standardised incidence rate was 20 per 100,000 person-years for men and 22 per 100,000 person-years for women.

#### 3.1. Incidence of CNS tumours

Age was a strong risk factor for CNS tumours. There was no clear effect of level of education on the incidence rates by age, for either men or women (Fig. 1), and in fact none of the socioeconomic and demographic variables evaluated in this study had a strong effect on the IRR. Nevertheless, small but significant increases in the IRR were observed for men in the agricultural class and for those in the highest income



Persons with unknown level of education not included

**Fig. 1** – Age-specific incidence rates per 100,000 person-years for CNS tumour by education amongst persons born in 1925–1973, diagnosed in Denmark, 1994–2003. Supplementary table shows the crude incidence rate and the incidence rate standardised by age (5-year age groups) and period (two 5-year periods) with the total study population as the standard and the incidence rate difference with basic school as the reference.

**Table 1 – Incidence rate ratios (IRRs) with 95% confidence intervals (95% CIs) for CNS tumours in Danish persons born in 1925–1973 and aged  $\geq 30$  years, by socioeconomic, demographic and health-related variables, Denmark, 1994–2003**

	Men			Women		
	Obs	IRR <sup>a</sup> (95% CI)	Adjusted IRR <sup>b</sup> (95% CI)	Obs	IRR <sup>a</sup> (95% CI)	Adjusted IRR <sup>b</sup> (95% CI)
<i>Level of education</i>						
Basic or high school	987	1.00	1.00	1341	1.00	1.00
Vocational education	1023	0.95 (0.87–1.04)	0.95 (0.87–1.03)	898	1.11 (1.02–1.21)	1.11 (1.02–1.21)
Higher education	495	1.00 (0.89–1.11)	0.96 (0.86–1.08)	534	1.09 (0.98–1.21)	1.09 (0.98–1.22)
Unknown	43	1.09 (0.81–1.48)	1.09 (0.80–1.47)	30	1.06 (0.73–1.52)	1.05 (0.73–1.52)
<i>Disposable income<sup>c</sup></i>						
Lowest (1st quartile)	652	1.03 (0.93–1.13)	1.02 (0.92–1.13)	763	1.03 (0.94–1.13)	1.05 (0.95–1.15)
Middle (2nd–3rd quartile)	1162	1.00	1.00	1310	1.00	1.00
Highest (4th quartile)	734	1.13 (1.03–1.24)	1.14 (1.03–1.25)	730	1.04 (0.95–1.14)	1.03 (0.93–1.13)
<i>Affiliation to work market<sup>d</sup></i>						
Working	1804	1.00	1.00	1634	1.00	1.00
Unemployed or other	229	0.91 (0.79–1.04)	0.92 (0.80–1.07)	417	0.98 (0.88–1.09)	1.00 (0.89–1.12)
Early retirement pensioner	173	1.03 (0.88–1.21)	1.05 (0.89–1.24)	337	1.05 (0.93–1.19)	1.10 (0.97–1.25)
<i>Social class<sup>e</sup></i>						
Creative core	180	1.18 (1.01–1.39)	1.17 (0.97–1.42)	55	1.01 (0.75–1.36)	0.94 (0.69–1.29)
Creative professional	422	1.04 (0.92–1.16)	1.03 (0.91–1.16)	260	1.06 (0.88–1.26)	0.99 (0.82–1.20)
Bohemian	14	1.02 (0.60–1.73)	1.01 (0.59–1.72)	9	1.25 (0.64–2.43)	1.19 (0.61–2.32)
Service	593	0.97 (0.88–1.07)	0.97 (0.87–1.07)	1585	1.00 (0.87–1.15)	0.96 (0.84–1.11)
Manual	990	1.00	1.00	231	1.00	1.00
Agricultural	164	1.24 (1.05–1.47)	1.23 (1.04–1.45)	64	1.27 (0.97–1.68)	1.27 (0.96–1.67)
Unknown	185	0.96 (0.82–1.12)	0.95 (0.81–1.11)	599	1.04 (0.89–1.21)	1.03 (0.88–1.20)
<i>Housing tenure</i>						
Owner-occupied	1814	1.00	1.00	1823	1.00	1.00
Rental	701	0.99 (0.91–1.08)	0.99 (0.91–1.09)	948	1.04 (0.96–1.13)	1.05 (0.97–1.14)
Unknown	33	0.88 (0.62–1.24)	0.88 (0.62–1.24)	32	1.04 (0.73–1.48)	1.05 (0.74–1.49)
<i>Size of dwelling (m<sup>2</sup>)</i>						
0–49	61	0.72 (0.56–0.94)	0.72 (0.56–0.94)	32	0.75 (0.53–1.07)	0.76 (0.53–1.08)
50–99	767	0.98 (0.89–1.08)	0.98 (0.90–1.08)	1025	1.02 (0.93–1.11)	1.02 (0.94–1.12)
100–149	1022	1.00	1.00	1086	1.00	1.00
$\geq 150$	698	1.05 (0.95–1.15)	1.04 (0.94–1.14)	660	1.00 (0.91–1.10)	0.99 (0.90–1.10)
<i>Cohabiting status</i>						
Married	1783	1.00	1.00	1840	1.00	1.00
Cohabiting	222	0.92 (0.80–1.07)	0.92 (0.80–1.06)	219	0.99 (0.86–1.15)	1.00 (0.87–1.16)
Single	298	1.02 (0.90–1.16)	1.02 (0.89–1.16)	174	0.93 (0.79–1.09)	0.93 (0.79–1.09)
Widow or widower	65	0.97 (0.75–1.24)	0.97 (0.76–1.25)	298	1.10 (0.97–1.26)	1.11 (0.97–1.26)
Divorced	180	0.90 (0.77–1.05)	0.91 (0.78–1.06)	272	0.96 (0.85–1.09)	0.96 (0.84–1.09)
<i>Type of district</i>						
Capital area	812	1.00	1.00	981	1.00	1.00
Provincial city	1295	0.94 (0.86–1.03)	0.95 (0.87–1.04)	1368	0.85 (0.79–0.93)	0.86 (0.79–0.93)
Rural area	318	0.96 (0.84–1.09)	0.97 (0.85–1.10)	315	0.86 (0.76–0.98)	0.87 (0.77–0.99)
Peripheral rural area <sup>f</sup>	123	0.84 (0.70–1.02)	0.85 (0.70–1.03)	139	0.86 (0.72–1.03)	0.87 (0.73–1.04)
<i>Ethnicity<sup>g</sup></i>						
Danish	2548	1.00	1.00	2803	1.00	1.00
Immigrant or descendant from western country	53	0.99 (0.76–1.31)	0.99 (0.75–1.30)	72	1.05 (0.83–1.33)	1.03 (0.81–1.30)
Immigrant or descendant from non-western country	68	0.90 (0.71–1.15)	0.90 (0.70–1.15)	78	1.05 (0.84–1.31)	1.00 (0.78–1.27)
<i>Charlson comorbidity index<sup>h</sup></i>						
None	2117	1.00	1.00	2471	1.00	1.00
1	291	1.17 (1.03–1.33)	1.18 (1.04–1.34)	202	0.96 (0.83–1.11)	0.96 (0.83–1.11)
$\geq 2$	140	1.19 (1.00–1.42)	1.20 (1.01–1.43)	130	1.12 (0.94–1.34)	1.13 (0.95–1.35)
<i>Depression</i>						
No	2505	1.00	1.00	2704	1.00	1.00
Yes	43	1.09 (0.80–1.47)	1.10 (0.81–1.48)	99	1.26 (1.03–1.55)	1.27 (1.04–1.56)

Table 1 – continued

	Men			Women		
	Obs	IRR <sup>a</sup> (95% CI)	Adjusted IRR <sup>b</sup> (95% CI)	Obs	IRR <sup>a</sup> (95% CI)	Adjusted IRR <sup>b</sup> (95% CI)
Schizophrenia or other psychosis						
No	2528	1.00	1.00	2774	1.00	1.00
Yes	20	0.77 (0.50–1.20)	0.78 (0.50–1.21)	29	0.88 (0.61–1.26)	0.89 (0.62–1.28)

a Adjusted for calendar period (in 5-year intervals) and age modelled as age and age<sup>2</sup> in years.

b Adjusted for calendar period and age (as above) and additionally for level of education and disposable income.

c Household income after taxation and interest, adjusted for number of persons in household; categorised by gender-specific distribution of household disposable income per person.

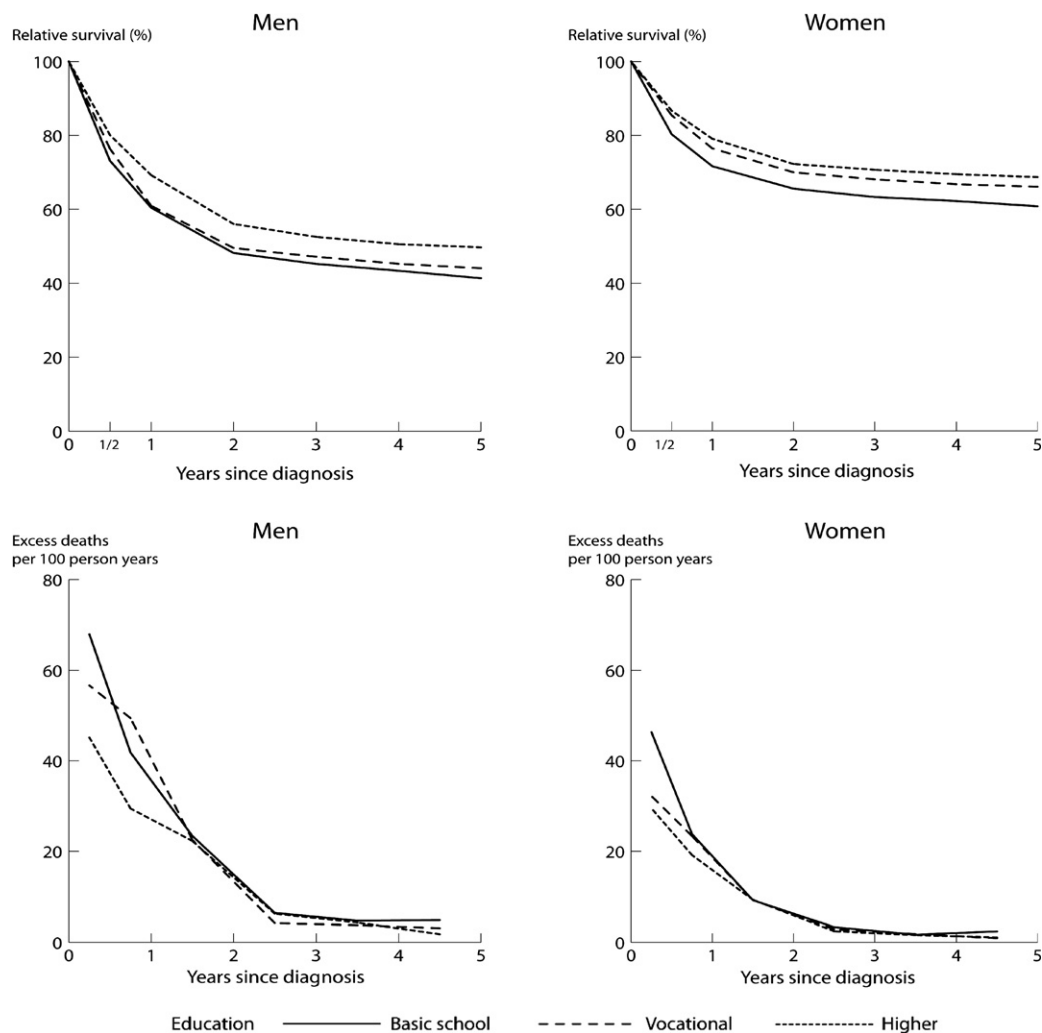
d For pensioners, work market affiliation before pension date was assigned and follow-up to age 69.

e Based on theory of creative class<sup>24</sup>: the creative core (e.g. researchers, designers, architects), creative professionals (e.g. managers, business and finance, lawyers, doctors), bohemians (e.g. artists, models), the service class (e.g. nurses, hairdressers, caterers), the manual class (e.g. construction workers, transport and production workers) and the agricultural class (e.g. farmers, fishermen).

f More than 40 km to a local centre with adequate possibilities for employment and not sharing a border with a centre municipality.

g Included as a separate indicator, but ethnic groups were excluded from the study population in all other analyses presented in this table, e.g. education and income.

h The presence of disorders, as defined in the Charlson index, was defined as an in- or outpatient contact with one of the diagnoses listed in Table 1 in<sup>12</sup> between 1978 and 2 years before the diagnosis of cancer. Grouped according to the accumulated sum of scores.



**Fig. 2 – Age-standardised relative survival and excess mortality rates per 100 person-years by level of education in patients born in 1925–1973, with CNS tumour diagnosed in Denmark 1994–2003 and followed through 2006. Relative survival is the ratio of the observed survival of the cancer patients and the survival that would have been expected if the patients had had the same age-, period- and education-specific mortality as the total study population. Excess mortality is excess to the same population mortality and estimated in intervals since diagnosis. Estimates were standardised for age on the basis of the age distribution of all patients with CNS tumour in the study cohort.**

**Table 2 – 1-year and 5-year relative survival (%) with 95% confidence interval (95% CI) by socioeconomic, demographic and health variables in patients aged  $\geq 30$  years born in 1925–1973, with CNS tumour diagnosed in Denmark between 1994 and 2003 and followed through 2006**

	Men					Women				
	No.	1-year survival		5-year survival		No.	1-year survival		5-year survival	
		%	95% CI	%	95% CI		%	95% CI	%	95% CI
Level of education										
Basic or high school	974	60	58–63	39	36–42	1336	72	69–74	58	55–61
Vocational education	1015	61	58–64	41	38–44	893	76	74–79	63	60–67
Higher education	494	69	65–74	47	43–52	532	79	75–83	66	62–71
Unknown	42	66	53–84	38	26–57	30	76	62–94	71	55–93
Disposable income <sup>b</sup>										
Lowest (1st quartile)	643	60	56–64	42	38–46	761	72	69–76	58	54–62
Middle (2nd–3rd quartile)	1152	61	58–64	40	37–43	1302	75	73–78	62	59–64
Highest (4th quartile)	730	67	63–70	43	39–47	728	76	72–80	65	61–70
Affiliation to work market <sup>c</sup>										
Working	1791	73	71–75	50	48–52	1630	83	81–85	69	67–72
Unemployed or other	228	71	64–78	45	38–54	415	84	80–88	71	66–76
Early retirement pensioner	168	59	51–69	39	31–51	332	75	69–82	64	56–73
Social class <sup>d</sup>										
Creative core	180	72	66–79	51	44–59	55	74	64–85	57	44–73
Creative professional	421	63	59–68	42	38–47	258	80	75–86	68	62–75
Bohemian	14	74	62–88	56	46–69	9	40	–	13	–
Service	590	65	62–69	43	39–47	1579	74	72–77	61	58–63
Manual	981	58	55–61	38	35–41	230	77	72–83	63	57–71
Agricultural	161	69	62–75	47	40–56	64	71	61–83	55	42–71
Unknown	178	57	50–65	38	30–48	596	72	68–76	61	56–66
Housing tenure										
Owner-occupied	1802	64	61–66	42	40–45	1813	75	73–77	62	60–64
Rental	691	59	55–62	39	35–43	946	74	71–77	60	57–64
Unknown	32	47	33–68	26	14–49	32	60	46–79	55	40–75
Size of dwelling (m <sup>2</sup> )										
0–49	60	51	40–65	39	29–54	31	62	48–80	56	41–76
50–99	756	60	56–63	38	35–42	1023	73	70–75	60	56–63
100–149	1018	62	59–65	41	38–44	1079	75	73–78	62	59–65
≥150	691	66	62–69	45	41–49	658	76	73–80	63	59–67
Cohabiting status										
Married	1773	63	61–65	41	39–44	1834	74	72–76	61	59–64
Cohabiting	221	59	52–68	41	34–50	218	78	71–85	70	62–78
Single	292	58	52–65	38	32–45	172	76	69–83	62	54–71
Widow/widower	63	66	55–79	51	41–63	297	71	62–81	57	46–71
Divorced	176	57	50–65	38	31–47	270	70	65–76	54	48–61
Type of district										
Capital area	806	61	58–65	39	36–43	980	75	72–78	61	58–65
Provincial city	1282	62	60–65	42	39–45	1358	74	72–76	62	60–65
Rural area	316	64	59–69	45	39–51	314	74	70–79	59	54–65
Peripheral rural area <sup>e</sup>	121	63	56–70	40	32–50	139	70	63–78	57	49–66
Ethnicity <sup>f</sup>										
Danish	2525	62	60–64	41	39–43	2791	74	73–76	61	59–63
Immigrant or descendant from western country	53	68	58–80	55	44–69	72	87	79–95	65	55–78
Immigrant or descendant from non-western country	68	70	56–87	47	35–63	78	85	77–93	72	62–84
Charlson comorbidity index <sup>g</sup>										
None	2102	63	61–65	42	40–44	2461	75	73–77	62	60–64
1	288	58	52–64	35	29–43	201	72	66–79	60	53–69
≥2	135	62	54–71	38	28–51	129	73	67–81	55	46–66

Table 2 – continued

	Men					Women				
	No.	1-year survival		5-year survival		No.	1-year survival		5-year survival	
		%	95% CI	%	95% CI		%	95% CI	%	95% CI
Depression										
No	2482	62	60–64	41	39–43	2694	74	73–76	61	59–63
Yes	43	70	57–85	53	39–74	97	74	67–83	63	54–75
Schizophrenia or other psychosis										
No	2505	62	60–64	41	39–43	2762	74	73–76	61	59–63
Yes	20	48	30–76	28	15–53	29	76	60–95	63	46–88

a Ratio of observed survival of CNS tumour patients and survival that would have been expected if the patients had had the same age-, period-, socioeconomic, demographic or health-related indicator-specific mortality as the total study population; for 'social class' and 'ethnicity', expected survival is adjusted only for age, not period, because of low power.

b Household income after taxation and interest, adjusted for number of persons in household; categorised by gender-specific distribution of household disposable income per person.

c For pensioners, work market affiliation before pension date was assigned and follow-up to age 69.

d Based on theory of creative class (24): the creative core (e.g. researchers, designers, architects), creative professionals (e.g. managers, business and finance, lawyers, doctors), bohemians (e.g. artists, models), the service class (e.g. nurses, hairdressers, caterers), the manual class (e.g. construction workers, transport and production workers) and the agricultural class (e.g. farmers, fishermen).

e More than 40 km to a local centre with adequate possibilities for employment and not sharing a border with a centre municipality.

f Excluded from the study population in all other analyses presented in this table.

g The presence of, as defined in the Charlson index, was defined as an in- or outpatient contact with one of the diagnoses listed in Table 1 in<sup>12</sup> between 1978 and 2 years before the diagnosis of cancer. Grouped according to the accumulated sum of scores.

quartile. Amongst women in the agricultural class the increased IRR was nonsignificant. A Charlson index score of 1 or more was associated with a higher IRR of CNS tumours in men, and a score of 2 or more was associated with a nonsignificant increase amongst women. The IRR amongst women was highest for those living in the capital area and those with diagnosed depression (Table 1). Diagnoses of schizophrenia and other psychotic diseases were associated with nonsignificantly decreased IRRs for CNS tumours in both sexes (Table 1).

An association between a previous diagnosis of chronic somatic disease and risk for CNS tumours was not apparent when only malignant tumours were analysed (data not shown).

### 3.2. Relative survival after CNS tumours

The 1-year relative survival after a CNS tumour was 62% for men and 74% for women, and that 5 years after diagnosis was 41% for men and 61% for women. The 1-year relative survival after diagnosis of a malignant tumour was 47% (95% CI, 45–50%) for men and 48% (95% CI, 45–51%) for women, and that after 5 years was 17% (95% CI, 15–19%) for men and 21% (95% CI, 19–24%) for women (data not shown).

Fig. 2 shows that the relative survival rates for men and women differed when both malignant and benign CNS tumours were included in the analyses. Nearly half the male patients died within the first 2 years, and most male patients died within the first 6 months after diagnosis. For women, the pattern was similar, but survival was generally longer. A difference in relative survival rate by educational level was seen for both the sexes after the first 6 months (Fig. 2). Fig. 2 also shows that the excess mortality rate was highest for men with basic schooling or vocational education in comparison with patients with higher education in the first 18

months after diagnosis. At later times, no clear difference by educational level was observed for men or women.

We observed a socioeconomic gradient in the age-standardised 1- and 5-year relative survival rates of both the sexes when measured by the social indicators education, size of dwelling and affiliation to the work market. A similar gradient with increasing relative survival rates with increasing social level was seen by social class, although for women only an effect in the 5-year relative survival rate was present. High income was associated with a higher relative survival rate although only significant for men after 1 year and women after 5 years (Table 2).

## 4. Discussion

Overall, we observed no major effects of socioeconomic position or demographic factors on the incidence of CNS tumours, in line with the previous reports from large Scandinavian register-based cohort studies,<sup>6,7,15</sup> as well as a large, recently published case-control study,<sup>16</sup> in which occupational title or educational level was used as the indicator of SEP.<sup>6,7,15</sup> A number of case-control studies have given conflicting results, with an increasing risk with increasing socioeconomic position when occupational title was used as the sole social indicator.<sup>3–5,17</sup> In a more recent interview-based case-control study, an increasing incidence of benign tumours but not malignant tumours was associated with increasing income and education.<sup>18</sup>

The increased IRR for CNS tumours observed amongst men assigned to the agricultural social class is in line with the results of several previous studies,<sup>19,20</sup> and is probably a result of occupational exposure rather than socioeconomic position. A high incidence of CNS tumours was observed amongst Danish farmers in the 1970s.<sup>6</sup>



Marital status has previously been shown to affect survival, with a slightly lower rate amongst single persons.<sup>8,21</sup> We did not observe any difference by demographic factors, apart from better survival of descendants of immigrants from non-western countries, based, however, on only small numbers.

Both short-term (1-year) and long-term (5-year) survival was better amongst women than men. This may reflect the fact that meningioma, which has a better prognosis than gliomas, is more frequent amongst women.<sup>1</sup> This conclusion is further supported by the observation that the large difference in survival between the sexes vanished when the analysis was restricted to malignant tumours.

We found that socioeconomic position, as assessed by level of education, size of dwelling, income, affiliation to the work market and amongst men additionally social class, is a prognostic factor for survival after a CNS tumour in Denmark. Our findings are in line with the results of the few previous studies that reported better survival after a CNS tumour in more socially advantaged groups. In a large cohort study in Finland in which social class were classified by occupation, social class IV had slightly poorer survival than social class I, but the result did not reach significance (RR, 0.79; 95% CI, 0.59–1.05).<sup>9</sup>

A similar pattern was observed in a cohort study in the United Kingdom of 30,489 patients with glioma, in which socioeconomic position was based on the Carstairs index.<sup>10,22</sup> The most affluent patients with CNS tumours had better survival than the most deprived patients, the effect being most pronounced during the first year after diagnosis.

In previous studies, social inequalities in cancer survival have been attributed partly to a greater burden of chronic somatic comorbidity in disadvantaged groups.<sup>23</sup> This has not to the best of our knowledge been evaluated by studying actual comorbidity in patients with CNS tumours. Excess somatic comorbidity cannot explain all the observed differences in the present study, as only decreased long-term survival in women was associated with the Charlson comorbidity index.

Better compliance with public health policies, including earlier contact with the health system resulting in earlier diagnosis, by patients with a higher socioeconomic position might explain the differences in relative survival rates by socioeconomic position; however, we had no information about the tumour size or disease stage at the time of diagnosis. Alternatively, the observed differences in survival after diagnosis of a CNS tumour might be due to differences in access to treatment, time to surgery and referral to or compliance with adjuvant treatment in our public health system.

### Conflict of interest statement

None declared.

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