Socioeconomic and geographic determinants of survival of patients with digestive cancer in France

O Dejardin*,1, L Remontet2, AM Bouvier3, A Danzon3, B Trétarre3, P Delafosse3, F Molinié3, N Maarouf3, M Veiten3, EA Saulaue4, N Bourdon-Raverdy3, P Grosclaude3, S Boutreux3, G De Pouvourville4 and G Launoy1,3

*Cancers & Populations’ ERI 3 INSERM; Faculty of Medicine, Avenue de la Côte de Nacre 14032 Coen Cedex, France; 2Biostatistic Analyses Unit of ‘Hospices Civils de Lyon’ (HCL), Lyon, France; 3French Network of Cancer Registries (FRANCIM), Toulouse, France; 4Centre for Research on Medicine, Science, Health and Society (CERMES/INSERM U750), Villejuif, France

Using a multilevel Cox model, the association between socioeconomic and geographical aggregate variables and survival was investigated in 81 268 patients with digestive tract cancer diagnosed in the years 1980–1997 and registered in 12 registries in the French Network of Cancer Registries. This association differed according to cancer site: it was clear for colon (relative risk (RR) = 1.10 (1.04–1.16), 1.10 (1.04–1.16) and 1.14 (1.05–1.23), respectively, for distances to nearest reference cancer care centre between 10 and 30, 30 and 50 and more than 90 km, in comparison with distance of less than 10 km; P-trend = 0.003) and rectal cancer (RR = 1.09 (1.03–1.15), RR = 1.08 (1.02–1.14) and RR = 1.12 (1.05–1.19), respectively, for distances between 10 and 30 km, 30 and 50 km and 50 and 70 km, P-trend = 0.024) (n = 28 010 and n = 18 080, respectively) but was not significant for gall bladder and biliary tract cancer (n = 2893) or small intestine cancer (n = 1038). Even though the influence of socioeconomic status on prognosis is modest compared to clinical prognostic factors such as histology or stage at diagnosis, socioeconomic deprivation and distance to nearest cancer centre need to be considered as potential survival predictors in digestive tract cancer.

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The incidence of digestive cancers is still increasing, mainly due to colorectal cancer trends (Parkin et al, 2002), and retains a poor prognosis, as reported by the European Cancer Registries (EUROCARE) study (Berrino et al, 2003). This study found a large range of digestive cancer survival within Europe, while a survival difference in colorectal cancer between Europe and the USA has also been reported (Ciccolallo et al, 2005). To explain such inter-country differences, besides the important effect of stage at diagnosis and surgery, the influence of health system organisation and social determinants needs to be further investigated.

The 5-year relative survival of the 60 000 cases of digestive cancer each year in France (Remontet et al, 2003) is reasonably high compared to the European mean. Nevertheless, a few French studies of colorectal cancer from a single cancer registry (Launoy et al, 1992; Desoubeaux et al, 1997) have suggested social inequalities in survival, underprivileged patients having poorer prognosis than more affluent patients. We recently suggested that access to specialised care was strongly associated with road distance to the nearest specialised centre especially for women and elderly patients (Dejardin et al, 2005). Recent research in different health-care systems also suggests a relationship between surgical volume and survival (Kee et al, 1999; Shrag et al, 2000; Platell et al, 2003). Geographical access to reference centres is therefore relevant to the study of social inequalities in cancer survival.

This study, based on data on all digestive cancers registered by the French Network of Cancer Registries (FRANCIM) between 1980 and 1997 in France, investigates the influence of the socioeconomic and geographical environment on survival of affected patients.

MATERIALS AND METHODS

Population and data

The study population comprised 81 268 patients with digestive tract cancer registered between 1 January 1980 and 31 December 1997 in 12 cancer registries (Calvados, Côte d’Or, Doubs, Hérault, Isère, Loire-Atlantique, Manche, Bas-Rhin, Haut-Rhin, Saône et Loire, Somme and Tarn), all belonging to the French Network of Cancer Registries (FRANCIM); as such, their exhaustiveness and data quality are regularly checked by the French Institute of Health and Medical Research (INSERM) and the International Agency for Research on Cancer (IARC). General characteristics of the study population, representing 14.4% of the French population (n = 8 441 366), are detailed in Table 1.

Survival was defined as the interval in months between diagnosis and last information on vital status. Two types of variable were used in the study, individual and aggregate.

*Correspondence: Dr O Dejardin; olivier.dejardin@unicaen.fr
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Table 1 Digestive cancer diagnosed between 1980 and 1997 in 12 French Communes

| Site                      | N   | %   |
|---------------------------|-----|-----|
| Oesophagus                | 781 3 | 9.61 |
| Stomach                   | 11 619 | 14.30 |
| Colon                     | 28 010 | 34.47 |
| Rectum                    | 18 080 | 22.25 |
| Small intestine bowel     | 1038 | 1.28 |
| Liver                     | 5797  | 7.13 |
| Gall bladder              | 2893 | 3.56 |
| Pancreas                  | 6018 | 7.41 |

Level 1: patient characteristics
- Age (years):
  - <60: 21 977 (29.77%)
  - 60–70: 11 910 (16.66%)
  - 70–78: 13 006 (16.00%)
  - >78: 32 155 (43.57%)
- Gender:
  - Men: 48 741 (59.98%)
  - Women: 32 527 (40.02%)

Year of diagnosis:
- 1980: 1653 (2.03%)
- 1981: 1690 (2.08%)
- 1982: 2284 (2.81%)
- 1983: 2424 (2.98%)
- 1984: 2450 (3.01%)
- 1985: 2436 (3.00%)
- 1986: 2519 (3.10%)
- 1987: 2554 (3.14%)
- 1988: 2516 (3.10%)
- 1989: 5400 (6.64%)
- 1990: 5541 (6.82%)
- 1991: 5990 (7.37%)
- 1992: 6331 (7.79%)
- 1993: 6405 (7.88%)
- 1994: 6976 (8.58%)
- 1995: 7892 (9.71%)
- 1996: 8039 (9.89%)
- 1997: 8168 (10.05%)

Level 2: commune characteristics
- Distance to nearest reference cancer care centre (km):
  - <10: 21 621 (26.60%)
  - 10–30: 13 924 (17.13%)
  - 30–50: 13 816 (17.00%)
  - 50–70: 12 569 (15.47%)
  - 70–90: 9 235 (11.36%)
  - >90: 10 103 (12.43%)

- Lost known status (at 1 January 2002):
  - Censored: 15 229 (18.74%)
  - Death: 61 820 (76.07%)
  - Lost of follow up: 4 243 (5.19%)

Level-2 aggregate variables: to compensate for the lack of individual information on socioeconomic status, each patient was attributed the socioeconomic characteristics of his neighbourhood (Commune level) using French Census data provided by the INSEE (1999) – National Institute for Statistics and Economic Studies, that is, proportion of people who did not complete secondary education to age 15, mean taxable income, unemployment rate, proportion of housing without bath or shower, manual worker proportion. Mean taxable income (2003) of each Commune was obtained from the ‘Direction Générale des Impôts’, the tax department.

French public health authorities have defined reference care centres as centres capable of providing care management of serious pathologies with bad prognosis as well as rare pathologies. In our study, the reference care centres are mainly composed of a university hospital and a specialised cancer centre. Road distance to nearest cancer reference care centre (always located in regional capitals) was calculated using MapInfo 6.5 (MapInfo corporation) and GIS (Geographical Information System). Road database (IGN route 500) of the National Geographic Institute (IGN) included 500 000 km of road. These variables were also considered as level-2 variables.

The population study was distributed within 5416 of the 6289 Communes of the 12 French counties (86.11%). The mean Commune population was 1342.1 (range 270 343 – 6). The mean number of cases per Commune was 15.00.

Analysis

A multilevel Cox proportional hazards model was used to take into consideration the hierarchical data structure (Goldstein, 1995). Level 1 included patient characteristics (age, sex and year of diagnosis), level 2 was composed of the Commune’s social indicator. All models were adjusted on department of residence (level 3).

Iterative general least square (IGLS) (Poisson first order marginal quasi likelihood (MQL)) was used to estimate the model. For each cancer site, we investigated the influence of each of six socioeconomic level-2 variables after adjustment on age, gender, year at diagnosis and department of residence in eight separate models. The influence of social variables on survival was investigated using linear trend for categorised variables.

Mlwin 2.01 (multilevel model project, London Institute of Education) with survival macro was used to perform the multilevel Cox model.

RESULTS

Table 2 shows the influence of socioeconomic and geographical environment on survival for patients with digestive tract cancer. The association between socioeconomic and geographical aggregate variables and survival differed by cancer site. This association was clear for colon and rectal cancer ($n = 28 010$ and $n = 18 080$, respectively) but not for gall bladder and biliary tract ($n = 2893$) or small intestine ($n = 1038$) cancers. For oesophageal, stomach, liver and pancreatic cancers, prognosis was associated with only one or two socioeconomic or geographic variables. Each of the socioeconomic aggregate variables was associated with at least one cancer site, the most frequently with distance to the nearest cancer reference care centre (hereafter ‘cancer centre’).
Table 2  Geographical and socioeconomic environment influence on survival for patient with digestive tract cancer (models adjusted for age, gender, year of diagnosis and department)

|                          | Oesophageal cancer (N = 7813) | Stomach cancer (N = 11619) | Small intestine cancer (N = 1038) | Colon cancer (N = 28010) |
|--------------------------|--------------------------------|-----------------------------|----------------------------------|--------------------------|
|                          | RR (95% CI)                     | RR (95% CI)                 | RR (95% CI)                      | RR (95% CI)              |
| Distance to nearest cancer care centre (km) |                          |                             |                                  |                          |
| <10                      | 1.00 (0.63–1.57)                 | 1.00 (0.65–1.58)             | 1.00 (0.63–1.57)                 | 1.00 (0.63–1.57)         |
| 10–30                    | 1.04 (0.70–1.56)                 | 1.06 (0.71–1.58)             | 1.08 (0.71–1.57)                 | 1.04 (0.70–1.56)         |
| 30–50                    | 1.03 (0.70–1.56)                 | 1.06 (0.71–1.56)             | 1.08 (0.71–1.56)                 | 1.04 (0.70–1.56)         |
| 50–70                    | 1.03 (0.70–1.56)                 | 1.06 (0.71–1.56)             | 1.08 (0.71–1.56)                 | 1.04 (0.70–1.56)         |
| 70–90                    | 1.04 (0.70–1.56)                 | 1.07 (0.71–1.57)             | 1.08 (0.71–1.57)                 | 1.04 (0.70–1.56)         |
| >90                      | 1.06 (0.70–1.57)                 | 1.09 (0.71–1.58)             | 1.09 (0.71–1.58)                 | 1.05 (0.70–1.57)         |
| Proportion who did not complete secondary education to age 15 (%) |                          |                             |                                  |                          |
| <14                     | 1.00 (0.63–1.57)                 | 1.00 (0.65–1.58)             | 1.00 (0.63–1.57)                 | 1.00 (0.63–1.57)         |
| 14–19                   | 1.03 (0.70–1.56)                 | 1.06 (0.71–1.56)             | 1.08 (0.71–1.56)                 | 1.04 (0.70–1.56)         |
| 19–23                   | 1.05 (0.70–1.56)                 | 1.07 (0.71–1.56)             | 1.08 (0.71–1.56)                 | 1.04 (0.70–1.56)         |
| >23                     | 1.04 (0.70–1.57)                 | 1.07 (0.71–1.57)             | 1.08 (0.71–1.57)                 | 1.04 (0.70–1.56)         |
| Mean taxable income (in euros) |                          |                             |                                  |                          |
| <13 458–14 997          | 1.00 (0.63–1.57)                 | 1.00 (0.65–1.58)             | 1.00 (0.63–1.57)                 | 1.00 (0.63–1.57)         |
| 14 997–16 711           | 1.03 (0.70–1.56)                 | 1.06 (0.71–1.56)             | 1.08 (0.71–1.56)                 | 1.04 (0.70–1.56)         |
| Unemployment rate (%)   | 1.00 (0.63–1.57)                 | 1.00 (0.65–1.58)             | 1.00 (0.63–1.57)                 | 1.00 (0.63–1.57)         |
| <7.9                    | 1.00 (0.63–1.57)                 | 1.00 (0.65–1.58)             | 1.00 (0.63–1.57)                 | 1.00 (0.63–1.57)         |
| 7.9–12                  | 0.96 (0.70–1.56)                 | 0.99 (0.71–1.56)             | 1.00 (0.71–1.56)                 | 1.00 (0.70–1.56)         |
| 12–15                   | 0.99 (0.70–1.56)                 | 1.02 (0.71–1.57)             | 1.03 (0.71–1.56)                 | 1.00 (0.70–1.56)         |
| >15                     | 1.02 (0.70–1.57)                 | 1.05 (0.71–1.58)             | 1.06 (0.71–1.57)                 | 1.05 (0.70–1.56)         |
| Proportion of houses without bath or shower (%) |                          |                             |                                  |                          |
| <1.1                    | 1.00 (0.63–1.57)                 | 1.00 (0.65–1.58)             | 1.00 (0.63–1.57)                 | 1.00 (0.63–1.57)         |
| 1.1–1.7                 | 1.06 (0.70–1.56)                 | 1.09 (0.71–1.56)             | 1.10 (0.71–1.56)                 | 1.06 (0.70–1.56)         |
| 1.7–2.7                 | 1.08 (0.70–1.56)                 | 1.10 (0.71–1.56)             | 1.10 (0.71–1.56)                 | 1.07 (0.70–1.56)         |
| >2.7                    | 1.12 (0.70–1.57)                 | 1.14 (0.71–1.58)             | 1.14 (0.71–1.57)                 | 1.07 (0.70–1.57)         |
| Proportion of manual workers (%) |                          |                             |                                  |                          |
| <13                     | 1.00 (0.63–1.57)                 | 1.00 (0.65–1.58)             | 1.00 (0.63–1.57)                 | 1.00 (0.63–1.57)         |
| 13–17                   | 0.93 (0.70–1.56)                 | 0.99 (0.71–1.56)             | 1.00 (0.71–1.56)                 | 1.02 (0.70–1.56)         |
| 17–21                   | 0.96 (0.70–1.56)                 | 0.99 (0.71–1.56)             | 1.00 (0.71–1.56)                 | 1.04 (0.70–1.56)         |
| >21                     | 0.93 (0.70–1.57)                 | 1.04 (0.71–1.58)             | 1.05 (0.71–1.57)                 | 1.03 (0.70–1.56)         |

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Oesophageal cancer

Distance to the nearest cancer centre was not associated with survival. The proportion of housing without bath or shower was significantly associated with prognosis (P-trend = 0.004). In comparison with patients living in a Commune with less than 1.1% houses without bath or shower, patients living in a Commune with a high proportion of housing without bath or shower had poorer prognosis (relative risk (RR) = 1.08 (1.00–1.16) and RR = 1.12 (1.04–1.20), respectively for 1.7–2.7% of housing without bath or shower).

Stomach cancer

Distance to nearest cancer centre was associated with prognosis (P-trend = 0.0014) even after adjustment for age, gender and department of residence, patients living far from a cancer centre having poorer prognosis than those living closer (RR = 1.09 (1.00–1.20) and RR = 1.11 (1.00–1.23), respectively, for 50–70, and >90 km).

The association of prognosis with the proportion of housing without bath or shower was close to significant (P = 0.08), being significant when the department of residence was excluded from the analysis; those in a Commune with a high proportion of housing without bath or shower having poorer prognosis (P-trend < 0.001) (results not shown).

Small intestine

Although the prognosis of cancer did not decrease with the distance to nearest cancer centre, the prognosis of patients living 30–50 km from a cancer centre was significantly poorer than those living less than 10 km away (RR = 1.48 (1.18–1.85)). An increasing proportion of manual workers was somewhat associated with poorer prognosis (P-trend = 0.072), becoming significant when the proportion of manual workers was used as a continuous variable (P-trend = 0.02) (results not shown).

Colon cancer

After adjustment for individual characteristics, patients living far from a cancer centre had poorer survival than those living closer (RR = 1.10 (1.04–1.16), 1.10 (1.04–1.16) and 1.14 (1.05–1.23), respectively, for distance of 10–30 km, 30–50 km and over 90 km, compared to less than 10 km, P = 0.003).

Patients living in a Commune with less than 14% of people who did not complete secondary education to age 15, patients living in a Commune with a greater proportion had a poorer survival (P-trend = 0.002). Patients living in a Commune with low mean taxable income (<13548 euros) had poorer survival than those in a more affluent Commune (RR = 0.92 (0.88–0.97) with mean taxable income >16711 euros) (P-trend = 0.002).

Rectum cancer

Patients living far from a cancer centre had poorer prognosis than those living closer (RR = 1.09 (1.03–1.15), RR = 1.08 (1.02–1.14) and RR = 1.12 (1.05–1.19), respectively, for distances to nearest cancer centre of 10–30, 30–50 and 50–70 km compared with less than 10 km, P-trend = 0.024). In comparison with patients living in a Commune with a low proportion of people who did not complete secondary education to age 15, those living in a Commune with a higher proportion of people who did not complete secondary education to age 15 had a poorer survival (RR = 1.07 (1.02–1.13) and RR = 1.09 (1.03–1.15), respectively, for proportion of people who did not complete secondary education to age 15 of 19–23 and >23%). In comparison with patients living in a Commune with less than 13% of manual workers, those living in a Commune with a higher proportion have poorer survival (RR = 1.07 (1.01–1.13), RR = 1.10 (1.04–1.16) and RR = 1.07 (1.01–1.14), respectively for proportion of manual workers between 13 and 17, 17 and 21 and >21%; P-trend = 0.009).
Liver

Distance to the nearest cancer centre was not associated with prognosis of liver cancer. Patients living in a Commune with a high unemployment rate had poorer survival than those in a less-deprived Commune (RR = 1.09 (1.01 – 1.18) and RR = 1.09 (1.00 – 1.19), respectively, for unemployment rate 12 – 15% and above 15%, compared with below 8%; P-trend = 0.010).

Gall bladder and Biliary tract

Neither socioeconomic variables nor distance to the nearest cancer centre were significantly associated with prognosis of gall bladder or biliary cancer.

Pancreas

Distance to the nearest cancer centre was not significantly associated with prognosis. Patients living in a Commune with a high proportion of manual workers had poorer prognosis than those living in a more affluent Commune (P-trend = 0.032). The association with unemployment rate was suggestive (P-trend = 0.073) and became significant (P = 0.034) when tested using a continuous variable.

DISCUSSION

These results support the hypothesis that socioeconomic and geographical environment influences prognosis of digestive cancer in France, as previously found in other European countries and in the USA (Woods et al, 2006). This influence appears to be significant for the most common cancers with better prognosis (colon, rectum), and to a lesser extent for cancers with poorer prognosis (oesophagus, stomach, liver and pancreas). No influence was found for gall bladder and biliary tract cancer.

Certain methodological aspects of our study are relevant. In the absence of individual-based social information, the surrogate of an aggregate (Commune level) social indicator was used. Moreover, no deprivation index is available in France, prompting a multidimensional approach to poverty (unemployed, income; working conditions and accommodation).

The level of geographical unit used in our study was the Commune. Communes are the smallest administrative unit in France but their population sizes vary widely. The National Institute for Statistics and Economic Studies (INSEE) has defined a smaller unit: ‘Ilots regroupés d’indice statistique’ (IRIS). Regional capital and other important Communes are divided into several IRIS and small Communes constitute one IRIS. Therefore, the use of IRIS increases the accuracy of socioeconomic variables in such studies (Woods et al, 2005). Unfortunately, the lack of information on patients’ addresses prevented us from using the IRIS in our study. Socioeconomic inequalities highlighted in our study are likely to have been increased had we used the IRIS unit.

The use of relative survival, instead of crude survival, would have enabled us to adjust for comorbidity associated with age and gender, but to our knowledge, no multilevel relative survival model is available. As recently stressed (Auvinen and Karjalainen, 1997), relative survival tends to underestimate social inequalities in survival so we have preferred multilevel analysis rather than specific mortality.

The present study suffers from the lack of information on change in Census data during the study period. In fact, aggregate variables were collected in 1999 by the French Census and in 2003 for income tax purposes in an annual survey by the Direction Générale des Impôts, the tax department. The National Institute for Statistics and Economic Studies does not organise a Census each year. Furthermore, since multiple comparisons were made (six factors were tested for each of the eight cancers), some tests (two or three) may be significant by chance.

Nevertheless, our results accord with previous studies that stressed the appreciable gap between the most deprived and the most affluent in survival (Woods et al, 2006). At least in England and Wales, this was more widespread in the 1990s than previously (Coleman et al, 2004). In France, two studies, based on small samples, demonstrated that Communes with a high proportion of dwellings without bath or shower were negatively correlated with cancer survival (Monnet et al, 1993) and that farmers of both sexes and unemployed men had poorer survival than other socioeconomic categories (Desoubeaux et al, 1997). The frequent association of this housing measure suggested that it was a relevant socioeconomic indicator.

For cancers with good prognosis, in our study, the influence of socioeconomic aggregate variables on survival were similar to those in a Norwegian population-based study also using multilevel survival (discrete-time hazard regression) (Kravdal, 2006). In the Norwegian study, education variables (at individual and community level) were the most relevant socioeconomic indicators.

In our study, unlike the Norwegian study (Kravdal, 2006), road distance between cancer centres and the Commune of residence is a predictor of cancer survival for almost half of the cancer sites. This contradiction between the two studies may be partially explained by the adjustment for cancer stage at diagnosis in the Norwegian study. Lack of information on stage at diagnosis in our study restricts the interpretation of our results. Cancer stage at diagnosis is often more advanced in deprived people (Ionescu et al, 1998; Ciconne et al, 2000), although one recent study (Brewster et al, 2001) failed to find this for colorectal cancer using Carstairs deprivation categories. The explanation of social inequalities in cancer survival by the difference in stage at diagnosis is attractive as a consequence of the importance of stage at diagnosis as a prognostic factor. Nevertheless, as underlined by a recent review (Woods et al, 2006), stage at diagnosis does not entirely account for the gap in cancer survival between the most deprived and the most affluent.

Although geographic and socioeconomic effects on prognosis are modest compared to clinical prognostic factors such as histology or stage at diagnosis, they warrant consideration as potential survival predictors in digestive tract cancer. Few such studies have been carried out, partly due to the lack of individual-based information. Our study implies that the French health-care system, which theoretically provides free access to care to all patients, with extensive hospital services throughout the country, does not prevent social inequalities in cancer management.

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