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Burden of gastric and digestive cancers in the French Caribbean: perspectives from population-based cancer registries of Martinique, Guadeloupe and French Guiana (2007-2014).

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TITLE PAGE

Burden of gastric and digestive cancers in the French Caribbean: perspectives from populationbased cancer registries of Martinique, Guadeloupe and French Guiana (2007-2014).

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Keywords: cancer registry, incidence, Caribbean, mortality. Digestive cancers

Strengths and limitations of this study

- The purpose of this project is to present incidence and mortality for digestive cancer in Guadeloupe, French Guiana and Martinique for the 2007-2014 period.
- This study will contribute to expanding knowledge on the epidemiology of world cancers with data from the Caribbean zone.
- Potential limitations include the fact that comorbidities and risk factors are not recorded and thus cannot be taken into account in statistical analyses.

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Burden of gastric and digestive cancers in the French Caribbean: perspectives from populationbased cancer registries of Martinique, Guadeloupe and French Guiana (2007-2014).

Abstract

Background

Data from population-based cancer registries contribute to improving our knowledge of Digestive cancer trends worldwide.

Objectives

In this study we present cancer incidence and mortality in Guadeloupe, French Guiana and Martinique for the periods 2008-2014, 2010-2014 and 2007-2014 respectively.

Design and Methods

Data were extracted from cancer registries. World-standardized incidence and mortality rates were calculated. Main digestive cancers were analysed, including oesophagus, stomach, colorectum, liver and pancreas cancers.

Results

We observed a global lower-incidence compared to mainland France, except for stomach cancer for which the incidence is high, with significant standardized incidence ratios (SIRs) in men and women at 1.90 vs 2.29 for Guadeloupe and French Guiana and 1.58 vs 2.31 for Martinique. We found a global lower-mortality, except for stomach cancer for which the mortality remains high, with significant mortality ratios (SMRs) in men and women at 2.10 vs 2.74 for Guadeloupe, 1.64 vs 1.79 for French Guiana and 2.05 vs 2.53 for Martinique. Overall, these 3 regions have similar world-standardized incidence (WSI) and mortality (WSM) rates which remain lower than those in mainland France. We noticed an overall over-incidence and over-mortality in men compared to women as in France.

Conclusions: there is a high incidence of stomach cancer in French overseas territories. Publication of these data contributed to expanding knowledge on the epidemiology of world cancers with data from the Caribbean zone.

Keywords: cancer registry, incidence, Caribbean, mortality. Digestive cancers

Introduction

A total number of 111 933 new cancer cases were estimated in the Caribbean in 2018 according to GLOBOCAN 2018 Database. The most common cancer types were prostate, breast, lung and colorectum cancers; cervical and stomach cancers had higher incidence rates compared to mainland France. Digestive cancer represented 20.6% of these incident cases in both sexes[1].

Main identified risk factors of digestive cancers include socio-economic status, chronic tobacco smoking, and alcoholism. Conversely, a diet rich in fruit and vegetables has been shown to have a protective effect[2]. Other risk factors of oesophageal adenocarcinoma include gastro-oesophageal reflux and obesity [3]. For stomach cancer, *Helicobacter pylori* infection, high intake of salt, exposure to N-nitroso compounds (through diet, tobacco and endogenous synthesis) have been identified as major causes of cancer development[4].

The French West-Indies have a particular socio-demographic profile compared to the Caribbean, with high life expectancy and favourable health indicators. Nevertheless, certain digestive cancers appear in over-incidence such as stomach cancer [4]and underline the need for a study of the evolution of cancers over time from the cancer registries. The epidemiological transition has begun and continues for these regions. At the same time, the development of the care offer provides innovative technical platforms.

The cancer control strategy implemented with the various cancer plans [5] has enabled the deployment of significant resources to reduce disparities in the face of cancer. Through the development of health promotion policies, the general population is made aware of the impact of certain risk factors on the development of cancer. Studies are therefore necessary in order to explore the evolution of digestive cancers in our regions [6-9].

Data from population-based cancer registries contribute to improving our knowledge of cancer trends worldwide. The cancer indicators generated are useful for the general population, for researchers, clinicians and local and governmental organisations, and to decision-makers in public health. Pooling of data from the three registries of the French overseas departments will make it possible to identify clinical and epidemiological characteristics of digestive cancers. In this study we present incidence and mortality for digestive cancer in Guadeloupe, French Guiana and Martinique for the 2007-2014 period.

Methods

Data sources for incidence and mortality

Incidence data

Data were extracted for Guadeloupe, French Guiana and Martinique for the periods 2008-2014, 2010-2014 and 2007-2014 respectively. They are coded according to the International Classification of Diseases for Oncology, Third Edition (ICD-O-3). For mainland France, incidence data (2007-2016 period) were estimated from healthcare and registry data, using a dedicated method described elsewhere [10].

Mortality data

Mortality data cover the period 2007 to 2014. All the data were extracted from the Centre for Epidemiology of the medical causes of death (CepiDC). Data for the year 2012 were not exploitable for Martinique.

Statistical methods

Standardized rates were calculated using the world standard population of the WHO as standard [12]. The standardized incidence ratio (SIR) or standardized mortality ratio (SMR) were calculated in this study, using incidence and mortality rates from mainland France as references. We present the average annual number of cases and deaths observed and the world-standardized incidence and mortality rates for these three regions, and at national level. The standardized incidence and mortality ratios for these three regions are also presented, with 95% confidence intervals.

Patient and Public Involvement

Our study did not involve direct patient contact. Patients were not involved in the design of this study. Regarding patient involvement, cancer cases are identified through multidisciplinary team meetings, through medical records and the registry, according to the Registry procedures (French National authority for the protection of privacy and personal data). Additional approval from ethical committees was not required.

Results

Main digestive cancers were analysed, including oesophagus, stomach, colorectum, liver and pancreas cancers. In our study, we observed a global lower-incidence in digestive cancer, except for stomach cancer.

Oesophagus

In Martinique, Guadeloupe and French Guiana, oesophageal cancer affects on average 34 men and 6 women per year (Table 1-2), i.e. 1.6% of incident cancer cases in men and 0.4% in women. It was

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Stomach

In the French West-Indies, 80 individuals were diagnosed with stomach cancer each year respectively in Guadeloupe and Martinique, and 20 in French Guiana. Stomach cancer is more common in these Departments than in mainland France, and represents 4.5% of cancers in men in Martinique, 5.3% in Guadeloupe and 5.7% in French Guiana.

In men, it is the 3rd most common malignancy in Martinique and Guadeloupe, and 4th most common in French Guiana. In women, it is the3rd most common malignancy in Martinique and 4th most common in Guadeloupe, whereas it is less common in French Guiana.

Observed incidence is comparable in men in Guadeloupe and French Guiana (respectively 14.3 and 14.6 per 100,000 person-years), and lower in Martinique (12.1). In women, world-standardized incidence rates are 6.9 in Martinique, 7.2 in French Guiana and 7.3 in Guadeloupe. This higher incidence of stomach cancer, with significant standardized incidence ratios (SIRs) at 1.90 for Guadeloupe and French Guiana and 1.58 for Martinique, place these 3 Departments at the top of the list of French regional incidence for this cancer.

In line with incidence data, mortality is also higher than in mainland France. In men, the worldstandardized mortality rate, which reached 9.3 per 100,000 person-years in Guadeloupe, 8.9 in Martinique and 7.3 in French Guiana, is between 1.6 and 2.1 times higher than the corresponding rate in mainland France (4.5) and in most French regions with the exception of Corsica. These differences are also evident for stomach cancer mortality in women, where the world-standard mortality rate of 4.4 per 100,000 person-years in Guadeloupe is more than 2.7 times higher than that of mainland France (1.7). Among the French West-Indies, Guadeloupe has the highest world-standard mortality rates, in both men and women.

Colon-Rectum

In the regions of Guadeloupe, Martinique and French Guiana, on average, 194 men and 173 women per year are diagnosed with CRC (Table 1-2), i.e. 9.0% of incident cancer cases in men and 12.2% in women. CRC was responsible for 78 deaths per year in men from 2007 to 2014, representing 9.1% of all cancer-related deaths, and 74 deaths per year in women (11.2%).

Overall, these 3 regions have similar world-standardized incidence rates, which remain lower than incidence in France as a whole; CRC mortality is also lower.

Liver

In Guadeloupe, Martinique and French Guiana, liver cancer is diagnosed in an average of 30 men and 14 women per year, accounting for 1.4% of incident cancer cases in men, and 1.0% in women. It was

responsible for 38 deaths per year in men from 2007 to 2014, representing 4.4% of cancer-related deaths, and 23 deaths per year in women (3.5% of cancer-related deaths).

Pancreas

In Martinique, Guadeloupe and French Guiana, pancreatic cancer is diagnosed in an average of 44 men and 39 women per year, accounting for 2.0% of incident cancers in men and 2.7% in women. It was responsible for 53 deaths per year between 2007 and 2014, corresponding to 6.2% of cancer deaths in men and 49 deaths per year (7.4% of cancer-related deaths) in women.

Discussion

Overall, Martinique, Guadeloupe and French Guiana present world-standardized incidence rates that vary somewhat between the three regions, but that are in general lower than overall rates for the whole of France; there is also lower mortality related to oesophageal cancer.

For the period 2007 to 2016, incidence of oesophageal cancer was 7.2 per 100,000 person-years in men, and 1.5 in women in mainland France, corresponding to a sex ratio of 4.85, and accounting for an average of 3,998 incident cases in men and 1,072 in women per year, i.e. 2.0% of incident cancer cases in men and 0.7% in women. Oesophageal cancer was responsible for 3,825 deaths per year in mainland France between 2007 and 2015, accounting for 3.4% of cancer-related deaths in men, and 1.3% in women.

Rates in France are around the average for Western European countries. The incidence of oesophageal cancer has been declining for several years in men, whereas it is increasing in women. Net survival at 5 years for the period 2005-2010 was 14% in men and 18% in women[11].

The majority of oesophageal cancers can be classed into two histological groups. Epidermoid cancers are the most frequent in France in both men and women [12]. Furthermore, the World Health Organization (WHO) has established that X-rays and gamma radiation can contribute to oesophageal cancer. The second histological type is adenocarcinoma, whose frequency looks set to exceed that of epidermoid oesophageal cancers in France beyond the year 2017 according to some projections[13]. The majority of adenocarcinomas of the oesophagus develop in the context of endobrachyoesophagus (Barrett's oesophagus), following the metaplasia-dysplasia-carcinoma sequence. The data used for this report do not make it possible to distinguish between these two histological types.

On average, 4,707 men and 2,587 women were diagnosed with stomach cancer each year over the period 2007-2016, accounting for 2.4% of incident cancer cases in men and 1.6% in women. Stomach cancer caused almost 4,600 deaths per year in mainland France over the period 2007-2014, i.e. 3.3% of cancer-related deaths in men, and 2.6% in women.

The estimated incidence of stomach cancer in France is amongst the lowest of all the estimates provided by the WHO [14]. Incidence of stomach cancer is on the decline over the long term, and this

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is confirmed by the observations between 2005 and 2012 [15]. Mortality has also been declining persistently since the 1990s[15].

Net survival at 5 years of patients diagnosed between 2005 and 2010 was estimated at 23% for men and 28% for women [11]. The risk of stomach cancer is increased in first-degree relatives of a patient with stomach cancer, in individuals who have undergone partial gastrectomy for cancer, or in those who have undergone endoscopic treatment for gastric cancer, as well as in case of precancerous lesions of the stomach and in persons originating from regions with high stomach cancer incidence. Obesity and gastro-oesophageal reflux are risk factors for cardia cancer [16-18]. In Guadeloupe, prevalence of Helicobacter pylori infection, the main risk factor for this type of cancer, was estimated to be 55% in blood donors. Further studies are required to estimate the prevalence in the general population, and among patients with cancer. Environmental risk factors, as well as high intake of salt, and smoked meat and fish in the French West Indies, as in certain Asian countries, could also contribute to the high incidence of stomach cancer. Consumption of fruit and vegetables has a protective effect against stomach cancer.

Colorectal cancer is also a cancer among those requiring a policy of care, from the stage of organized or individual cancer screening to follow-up as part of the various treatments delivered. Several studies have been carried out in Martinique by the Martinique Cancer Registry, on the evolution of this cancer but also on the factors conditioning survival by age at diagnosis [6 8 9]. Additional studies are underway to analyze regional survival within the French West-Indies and should allow a better understanding of the profile of cancer patients in our region. In France, this cancer benefits from a prevention program which involves health actors from the general practitioner to integration within the framework of the hospital care pathway. Over the period 2007 to 2016, on average, 22,828 men and 19,174 women were diagnosed each year in mainland France, accounting for 12% of incident cancer cases in men and women.

On average, over the period 2007-2016, 6,989 men were diagnosed with liver cancer each year, corresponding to 3.5% of incident cancer cases in men. The number of deaths from liver cancer annually was on average 5,739 for the period 2007-2014 in mainland France in men (i.e. 6.4% of all cancer-related deaths in men) and 2,118 in women (3.3% of all cancer-related deaths in women).

Liver cancer includes primary carcinoma of the liver or hepatocellular carcinoma (HCC, which represents more than 80% of all liver cancers), as well as cancer of the intra-hepatic bile ducts. It is more frequent in France than in Europe as whole or in developed countries [14].

In men, mortality has been declining since 1995, but the opposite trend has been observed in women [15]. For recently diagnosed cases (2005-2010), net survival at 5 years was 15% [19].

There is also under-mortality from liver cancer in men. Conversely, in women, world-standardized mortality rates show no significant excess- or under-mortality compared to the rates observed in mainland France.

There are numerous established risk factors for primary liver cancer [20] namely: alcohol consumption, hepatitis B and C viruses, obesity [20], diabetes and tobacco smoking. Bile duct cancer risk factors include liver fluke, biliary tract diseases (primary sclerosing cholangitis and Caroli disease), and exposure to certain chemical compounds such as dichloropropane or dichloromethane[21].

Geographic variations in incidence observed over the study period could be explained by heterogeneity in the prevalence of chronic diseases linked to alcohol use, hepatitis B and C, or steatosis [22]. Progress in the management of patients with cirrhosis enables carcinogenesis to continue, and also contributes to the increased number of liver cancer cases[22], notably cancer of the intrahepatic bile ducts, whereas other histological types, such as HCC, are on the decline[23]. The distinct geographic distribution of the main risk factors for liver cancer, particularly alcohol consumption, likely explains to a large extent the disparities between Departments in terms of incidence and mortality.

On average, 5,581 men and 5,278 women were diagnosed with cancer of the pancreas each year in mainland France over the period 2007-2016, corresponding to 2.8% of incident cancers in men and 3.3% in women. Pancreatic cancer caused 9,409 deaths per year in mainland France from 2007 to 2014, i.e. 5.4% of cancer-related deaths in men, and 7.3% in women.

The incidence of pancreatic cancer is the highest in Eastern Europe, France and Japan[24]. In France, the incidence rate for pancreatic cancer is higher than the average of the 28 EU countries [15]. Pancreatic cancer is one of the 10 most frequent types of cancer, and its incidence increased in both sexes between 2005 and 2012, whereas mortality has remained practically unchanged since the 1980s[15]

During the period 2005-2010, net survival at 5 years was 8% in men and 7% in women[11]. Overall, the 3 regions have world-standardized incidence rates that remain lower than those of France as a whole; mortality from pancreatic cancer is also lower.

Tobacco smoking and obesity are established environmental risk factors for pancreatic cancer. Lowcalorie diets, high alcohol consumption and increased abdominal fat have also been reported to be precipitating factors, while a diet rich in fruit, vegetables and folates, and regular physical exercise are reported to have a protective effect [22].

Conclusion

 Collaborative projects to promulgate this expertise will help to improve knowledge of the clinical, demographic, socio-economic or organizational factors that contribute to the heterogeneity of cancer pathologies in the region.

The next projects of analysis will allow the carrying out of cancer mapping studies as well as the implementation of studies on the risk factors of stomach cancer and the prevalence of Helicobacter Pylori infection in the Antilles.

The study of environmental and behavioral factors is therefore an important issue for a better understanding of the determinants of health and cancer survival.

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DECLARATIONS

Authors' contributions

LID, SB, CJ, JVB, JD were major contributors in writing the manuscript, made substantial contributions to conception and design, JP, BBM, JPe, EC, JM, MB revising it critically for important intellectual content. EC and FRANCIM Network made substantial contributions to conception and design; and revising it critically for important intellectual content. All authors read and approved the final manuscript.

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Conflict interest statement

The authors declare that there are no conflicts of interest.

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Data Sharing

Requests to access the data are welcome and will be considered by the Scientific Committee. For more information, please contact Clarisse Joachim, at <u>Clarisse.joachim@chu-martinique.fr</u>

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			In	cidence				- 1	Mortali	ity		
	Ne	ew cases ¹		WSR ²		SIR ³			TI D	WSR ²		SMR ³
Oesophagus												
Guadeloupe	18	[15;21]	6.1	[5.1;7.4]	0.78	[0.65; 0.93]	14	[12;17]	ž 4.8	[4.0;5.9]	0.89	[0.73 ; 1.07
Martinique	12	[10;15]	3.8	[3.1;4.8]	0.52	[0.42; 0.63]	10	[7;12]	ð 2.8	[2.2;3.8]	0.56	[0.43; 0.71
French Guiana	4	[2;6]	4.5	[2.7;7.2]	0.56	[0.34 ; 0.88]	2	[1;4]	^S 2.8	[1.6;4.6]	0.52	[0.31; 0.82
Mainland France	3998	[3851 ; 4153]	7.23	[6.95 ; 7.51]			3031	[2993 ; 3070]	5.18	[5.11; 5.25]		
Stomach									vnl			
Guadeloupe	50	[44 ; 55]	14.3	[12.8;16.1]	1.90	[1.70; 2.11]	32	[29;37]	9.3	[8.2;10.7]	2.10	[1.85; 2.37
Martinique	43	[39;48]	12.1	[10.8;13.7]	1.58	[1.42; 1.76]	33	[29;38]	8.9	[7.7;10.3]	2.05	[1.79; 2.33
French Guiana	14	[11;17]	14.6	[11.2; 18.7]	1.90	[1.48;2.41]	6	[5;8]	5 7.3	[5.3 ; 9.8]	1.64	[1.21; 2.17
Mainland France	4707	[4561 ; 4859]	7.88	[7.62; 8.14]			2952	[2914;2990]	4.54	[4.48;4.61]		
Colon-Rectum								ť]			
Guadeloupe	81	[74 ; 88]	26.2	[24.0 ; 28.6]	0.66	[0.61; 0.72]	34	[30;39]	10.1 10.8 6.3	[8.9; 11.5]	0.74	[0.66 ; 0.84
Martinique	90	[83;97]	26.9	[24.8;29.2]	0.70	[0.65; 0.76]	39	[35;44]	10.8	[9.5;12.4]	0.80	[0.71; 0.90
French Guiana	23	[19;27]	25.0	[20.4;30.3]	0.70	[0.57; 0.84]	5	[3;6]	6.3	[4.4 ; 8.7]	0.46	[0.33; 0.64
Mainland France	22828	[22442 ; 23222]	37.8	[37.2;38.5]			8976	[8910; 9041]	12.7	[12.6; 12.8]		-
Liver						N.			<u>.</u>			
Guadeloupe	10	[8;13]	3.6	[2.8;4.7]	0.27	[0.21; 0.33]	15	[12;18]	4.8	[3.9;5.8]	0.50	[0.41 ; 0.59
Martinique	11	[9;13]	3.3	[2.6; 4.3]	0.27	[0.22; 0.33]	18		g 5.4	[4.5;6.7]	0.57	0.48; 0.68
French Guiana	9	[7;12]	10.5	[7.6; 14.3]	0.85	[0.62; 1.13]	5		§ 5.9	[4.2;8.2]	0.66	0.47; 0.91
Mainland France		[6730 ; 7261]	12.38	[11.91; 12.87]				[5686 ; 5791]	9.25	[9.16; 9.34]		2
Pancreas									22			
Guadeloupe	15	[13;19]	4.8	[3.9;6.0]	0.51	[0.42; 0.62]	24	[20;27]	g 7.0	[6.0; 8.2]	0.94	[0.81 ; 1.08
Martinique	24	[20;27]	7.0	[6.0; 8.2]	0.75	[0.65; 0.87]	25	[22;29]	6.9	[5.8; 8.2]	0.94	0.80; 1.09
French Guiana		[3;7]	5.5	[3.4;8.3]	0.58	[0.37; 0.86]	4	[22 ; 29] [3 ; 6] [4760 ; 4857]	<u></u> 5.8	[4.0; 8.3]	0.69	0.48;0.96
Mainland France		[5460; 5705]	9.52	[9.30; 9.74]			4808	[4760 ; 4857]	7.70	[7.62; 7.79]		-

BMJ Open Table 1.Annual number of new cases and deaths for digestive cancers in men, standardized incidence and mortality rated, standardized incidence and mortality

(1) Incidence mainland France: 2007-2016 ; Guadeloupe : 2008-2014 ; Martinique : 2007-2014 ; French Guiana: 2010-2014. (2) World-standardized rates: rates are standardized to the age structure of the world standard population and expressed per 100,000 person-years. (3) Ratios standardized to mainland ected by copyright. France.

	Incidence									ty		
	New cases ¹		WSR ² SI			SIR ³	IR ³ Deaths			WSR ²	SMR ³	
Oesophagus									orua			
Guadeloupe	3	[2.0;5.0]	0.9	[0.5;1.5]	0.52	[0.33; 0.79]	2	[1;3]	₹ 0.4	[0.2;0.9]	0.45	[0.25; 0.75
Martinique	2	[2.0;4.0]	0.6	[0.3;1.1]	0.40	[0.24; 0.61]	2	[1;4]	202 0.6	[0.3;1.2]	0.55	[0.32; 0.88
French Guiana	1	[0.0;3.0]	1.2	[0.4;2.8]	0.82	[0.30; 1.79]	0	[0;1]	N 0.4	[0.1;1.4]	0.43	[0.09; 1.27
Mainland France	1072	[1018 ; 1130]	1.49	[1.41; 1.58]			794	[774;813]	∂ 0.96	[0.93 ; 0.98]		
Stomach									wnlo			
Guadeloupe	32	[28;37]	7.3	[6.3;8.6]	2.29	[2.00; 2.61]	23	[19;26]	ad 4.4	[3.7;5.3]	2.74	[2.35;3.10
Martinique	34	[30 ; 39]	6.9	[6.0 ; 8.0]	2.31	[2.04;2.60]	22		<u>8</u> 3.7	[3.1;4.7]	2.53	[2.15; 2.94
French Guiana	8	[5;10]	7.2	[5.0; 10.1]	2.29	[1.62; 3.15]	3	[2;4]	ਰੂ 3.3	[2.1;5.1]	1.79	[1.15; 2.6
Mainland France	2587	[2508;2670]	3.22	[3.11; 3.34]			1665	[1637; 1694]	∃1.71	[1.68; 1.75]		
Colon-Rectum									ttp:			
Guadeloupe	70	[64;76]	17.3	[15.7; 19.2]	0.68	[0.62; 0.75]	29	[26;33]	6 .2	[5.4;7.3]	0.76	[0.66; 0.8
Martinique	85	[79;92]	20.4	[18.8;22.3]	0.80	[0.74 ; 0.86]	41	[36;46]	6.2 7.8	[6.8 ; 9.0]	0.98	[0.87; 1.1
French Guiana	18	[14;22]	17.4	[13.8;21.7]	0.75	[0.60; 0.92]	4	[3;6]	8 3.8	[2.5; 5.5]	0.54	[0.37; 0.7
Mainland France	19174	[18895 ; 19458]	24.4	[24.0;24.8]		VIA	7938	[7877;8000]	g 7.5	[7.4;7.6]		
Liver						· N	•		nj.c			
Guadeloupe	4	[3;6]	1.1	[0.7;1.8]	0.44	[0.30; 0.63]	9	[7;11]	2 .0	[1.5; 2.7]	0.84	[0.65;1.0
Martinique	7	[5;9]	1.8	[1.3;2.7]	0.64	[0.48 ; 0.84]	12	[9 ; 15]	g 2.5	[1.9; 3.4]	1.04	[0.83; 1.2
French Guiana	3	[2;5]	2.7	[1.5;4.8]	1.21	[0.66 ; 2.02]	2		≥ 1.8	[1.0; 3.2]	0.86	[0.47; 1.4]
Mainland France		Not Available					2118	[2087;2151]	≝ 2.24	[2.20; 2.29]		-
Pancreas									23,			
Guadeloupe	12	[9;15]	3.0	[2.3;3.9]	0.43	[0.34 ; 0.53]	22	[19 ; 26]	202 4.8	[4.0;5.7]	0.98	[0.84;1.1
Martinique	22	[19;25]	4.4	[3.7;5.4]	0.75	[0.64 ; 0.87]	24	[21;28]	<u>₽</u> 4.4	[3.6;5.3]	0.97	[0.83;1.1
French Guiana	5	[3;7]	4.8	[3.0;7.4]	0.76	[0.48;1.13]	3	[2;5]	<u>S</u> 3.2	[2.0;5.0]	0.72	[0.47;1.0
Mainland France	5278	[5152; 5407]	6.46	[6.29; 6.63]		-	4601	[4554 ; 4648]	ត 4.83	[4.77 ; 4.89]		

BMJ Open Table 2. Annual number of new cases and deaths for digestive cancers in women, standardized incidence and mortality ates, standardized incidence and

(1) Incidence mainland France: 2007-2016 ; Guadeloupe : 2008-2014 ; Martinique : 2007-2014 ; French Guiana: 2010-2014. (2) World-standardized rates: rates are standardized to the age structure of the world standard population and expressed per 100,000 person-years. (3) Ratios standardized to mainland ected by copyright. France.

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STROBE Statement

Pa	ge 17 of 18		BMJ Open BMJ Open	
1 2			STROBE Statement Stratement	
3 4	Section/Topic	Item No	Recommendation 47	Reported on Page No
5 6 7	Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract 9 (b) Provide in the abstract an informative and balanced summary of what was done and what was found 8 9	1
8	Introduction			
9 10	Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	2
11	Objectives	3	State specific objectives, including any prespecified hypotheses	2
12	vietnoas			
13	Study design	4	Present key elements of study design early in the paper	3
15 16	Satting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up and data collection	3
17 18 19 20 21 22 23 24 25	Participants	6	 (a) Cohort study—Give the eligibility criteria, and the sources and methods of selection of participants. Bescribe methods of follow-up Case-control study—Give the eligibility criteria, and the sources and methods of case ascertainment and control selection. Give the rationale for the choice of cases and controls Cross-sectional study—Give the eligibility criteria, and the sources and methods of selection of participants. (b) Cohort study—For matched studies, give matching criteria and number of exposed and unexposed Case-control study—For matched studies, give matching criteria and the number of controls per case 	3
26 27 28	Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if	3
29 30		8*	For each variable of interest, give sources of data and details of methods of assessment (measurement). $\mathbf{\underline{\underline{B}}}$ escribe comparability of assessment methods if there is more than one group $\mathbf{\underline{B}}$	3
31 32	Bias	9	Describe any efforts to address potential sources of bias	3
	Study size	10	Explain how the study size was arrived at	3
34		11	Explain how quantitative variables were handled in the analyses. If applicable, describe which grouping servere chosen and why	3
35			(a) Describe all statistical methods, including those used to control for confounding	3
36 37			(b) Describe any methods used to examine subgroups and interactions	3
38			(c) Explain how missing data were addressed	3
39 40	Statistical methods	12	(d) Cohort study—If applicable, explain how loss to follow-up was addressed	3
40 41			<i>Case-control study</i> —If applicable, explain how matching of cases and controls was addressed	
42			<i>Cross-sectional study</i> —If applicable, describe analytical methods taking account of sampling strategy (e) Describe any sensitivity analyses	
43			(e) Describe any sensitivity analyses	Not
44 45			For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml	1

		BMJ Open	Page 18 of 1
		ор ел - 2	applicable
Section/Topic	Item No	Recommendation 71	Reported on Page No
Results		9 N	
Dentisiaente	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for aligibility, confirmed eligible, included in the study, completing follow-up, and analysed	3-5
Participants	13*	(b) Give reasons for non-participation at each stage	3-5
		(c) Consider use of a flow diagram	3-5
	144	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders	3-5
Descriptive data	14*	(b) Indicate number of participants with missing data for each variable of interest	3-5
		(c) Cohort study—Summarise follow-up time (eg, average and total amount)	3-5
		Cohort study—Report numbers of outcome events or summary measures over time	3-5
Outcome data	15*	Case-control study—Report numbers in each exposure category, or summary measures of exposure	3-5
		Cross-sectional study—Report numbers of outcome events or summary measures	3-5
		(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 5% confidence interval).	3-5
Main results	16	Make clear which confounders were adjusted for and why they were included	
Main results	16	(b) Report category boundaries when continuous variables were categorized	3-5
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	3-5
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	Not applicable
Discussion			
Key results	18	Summarise key results with reference to study objectives	5-7
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	5-7
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	5-7
Generalisability	21	Discuss the generalisability (external validity) of the study results	5-7
Other Information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	8
*Give information separately	for cases	and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross ectional studies.	
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Burden of gastric and digestive cancers in the French Caribbean: perspectives from population-based cancer registries of Martinique, Guadeloupe and French Guiana (2007-2014).

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Primary Subject Heading :	Epidemiology
Secondary Subject Heading:	Oncology
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Burden of gastric and digestive cancers in the French Caribbean: perspectives from populationbased cancer registries of Martinique, Guadeloupe and French Guiana (2007-2014).

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Keywords: cancer registry, incidence, Caribbean, mortality. Digestive cancers

or per review only

Burden of gastric and digestive cancers in the French Caribbean: perspectives from populationbased cancer registries of Martinique, Guadeloupe and French Guiana (2007-2014).

Abstract

Objectives: Data from population-based cancer registries contribute to improving our knowledge of Digestive cancer trends worldwide. In this study we present cancer incidence and mortality in Guadeloupe, French Guiana and Martinique for the periods 2008-2014, 2010-2014 and 2007-2014 respectively.

Design: Data were extracted from population-based cancer registries. World-standardized incidence and mortality rates were calculated. Main digestive cancers were analysed, including oesophagus, stomach, colorectum, liver and pancreas cancers.

Setting: This study was performed based on data from French Territories in the Caribbean Results:

We observed a lower-incidence compared to mainland France, except for stomach cancer for which the incidence is high, with significant standardized incidence ratios (SIRs) in men and women at 1.90 vs 2.29 for Guadeloupe and French Guiana and 1.58 vs 2.31 for Martinique. We found a lower-mortality, except for stomach cancer for which the mortality remains high, with significant mortality ratios (SMRs) in men and women at 2.10 vs 2.74 for Guadeloupe, 1.64 vs 1.79 for French Guiana and 2.05 vs 2.53 for Martinique. Overall, these 3 regions have similar world-standardized incidence (WSI) and mortality (WSM) rates which remain lower than those in mainland France. We noticed an overall high incidence and high mortality in men compared to women as in France.

Conclusions: There is a high incidence of stomach cancer in French overseas territories. Publication of these data contributed to expanding knowledge on the epidemiology of world cancers with data from the Caribbean zone.

Keywords: cancer registry, incidence, Caribbean, mortality. Digestive cancers **Strengths and limitations of this study**

- The purpose of this project is to present incidence and mortality for digestive cancer in Guadeloupe, French Guiana and Martinique for the 2007-2014 period.
- This study will contribute to expanding knowledge on the epidemiology of world cancers with data from the Caribbean zone.
- Potential limitations include the fact that comorbidities and risk factors are not recorded and thus cannot be taken into account in statistical analyses.

Introduction

A total number of 111 933 new cancer cases were estimated in the Caribbean in 2018 according to GLOBOCAN Database. The most common cancer types were prostate, breast, lung and colorectum cancers; cervical and stomach cancers had higher incidence rates compared to mainland France. Digestive cancer represented 20.6% of these incident cases in both sexes[1].

Main identified risk factors of digestive cancers include socio-economic status, chronic tobacco smoking, and alcoholism. Conversely, a diet rich in fruit and vegetables has been shown to have a protective effect[2]. Other risk factors of oesophageal adenocarcinoma include gastro-oesophageal reflux and obesity [3]. For stomach cancer, *Helicobacter pylori* infection, high intake of salt, exposure to N-nitroso compounds (through diet, tobacco and endogenous synthesis) have been identified as major causes of cancer development[4].

The French West-Indies have a particular socio-demographic profile compared to the Caribbean, with high life expectancy and favourable health indicators. Nevertheless, certain digestive cancers appear in over-incidence such as stomach cancer [4] and underline the need for a study of the evolution of cancers over time from the cancer registries.

The cancer control strategy implemented with the various cancer plans in France [5] has enabled the deployment of significant resources to reduce disparities in the face of cancer. Through the development of health promotion policies, the general population is made aware of the impact of certain risk factors on the development of cancer. Studies are therefore necessary in order to explore the evolution of digestive cancers in our regions [6-9].

Data from population-based cancer registries contribute to improving our knowledge of cancer trends worldwide. The cancer indicators generated are useful for the general population, for researchers, clinicians and local and governmental organisations, and to decision-makers in public health. Pooling of data from the three registries of the French overseas departments will make it possible to identify clinical and epidemiological characteristics of digestive cancers. In this study we present incidence and mortality for digestive cancer in Guadeloupe, French Guiana and Martinique for the 2008-2014, 2010-2014 and 2007-2014 periods respectively.

Methods

Data sources for incidence and mortality

Incidence data

Data were extracted for Guadeloupe, French Guiana and Martinique for the periods 2008-2014, 2010-2014 and 2007-2014 respectively. They are coded according to the International Classification of Diseases for Oncology, Third Edition (ICD-O-3). For mainland France, incidence data (2007-2016 period) were estimated from healthcare and registry data, using a dedicated method described elsewhere [10].

Mortality data

Mortality data cover the period 2007 to 2014. All the data were extracted from the Centre for Epidemiology of the medical causes of death (CepiDC). Data for the year 2012 were not exploitable for Martinique.

Statistical methods

Standardized rates were calculated using the world standard population of the WHO as standard [11]. The standardized incidence ratio (SIR) or standardized mortality ratio (SMR) were calculated in this study, using incidence and mortality rates from mainland France as references. We present the average annual number of cases and deaths observed and the world-standardized incidence and mortality rates by regions, and at national level. The standardized incidence and mortality ratios for these three regions are also presented, with 95% confidence intervals.

Patient and Public Involvement

Our study did not involve direct patient contact. Patients were not involved in the design of this study. Regarding patient involvement, cancer cases are identified through multidisciplinary team meetings, through medical records and the registry, according to the Registry procedures (French National authority for the protection of privacy and personal data). Additional approval from ethical committees was not required.

Results

Main digestive cancers were analysed, including oesophagus, stomach, colorectum, liver and pancreas cancers. In our study, we observed a lower-incidence in all digestive cancer, except for stomach cancer.

Oesophagus

In Martinique, Guadeloupe and French Guiana, oesophageal cancer affects on average 34 men and 6 women per year (Table 1-2), i.e. 1.6% of incident cancer cases in men and 0.4% in women. It was

responsible for 26 deaths per year in men between 2007 and 2014 (Table 1), representing 3.0% of cancer deaths, and 4 deaths per year in women (0.6%).

Stomach

In the French West-Indies, 80 individuals were diagnosed with stomach cancer each year respectively in Guadeloupe and Martinique, and 20 in French Guiana. Stomach cancer is more common in these Departments than in mainland France, and represents 4.5% of cancers in men in Martinique, 5.3% in Guadeloupe and 5.7% in French Guiana.

In men, it is the 3rd most common malignancy in Martinique and Guadeloupe, and 4th most common in French Guiana. In women, it is the 3rd most common malignancy in Martinique and 4th most common in Guadeloupe, whereas it is 8th in French Guiana.

Observed incidence is comparable in men in Guadeloupe and French Guiana (respectively 14.3 and 14.6 per 100,000 person-years), and lower in Martinique (12.1). In women, world-standardized incidence rates are 6.9 in Martinique, 7.2 in French Guiana and 7.3 in Guadeloupe. This higher incidence of stomach cancer, with significant standardized incidence ratios (SIRs) at 1.90 for Guadeloupe and French Guiana and 1.58 for Martinique, place these 3 Departments at the top 3 of French regional incidence for this cancer.

In line with incidence data, mortality is also higher than in mainland France. In men, the worldstandardized mortality rate, which reached 9.3 per 100,000 person-years in Guadeloupe, 8.9 in Martinique and 7.3 in French Guiana, is between 1.6 and 2.1 times higher than the corresponding rate in mainland France (4.5) and in most French regions with the exception of Corsica. These differences are also evident for stomach cancer mortality in women, where the world-standard mortality rate of 4.4 per 100,000 person-years in Guadeloupe is more than 2.7 times higher than that of mainland France (1.7). Among the French West-Indies, Guadeloupe has the highest world-standard mortality rates, in both men and women.

Colon-Rectum

In the regions of Guadeloupe, Martinique and French Guiana, on average, 194 men and 173 women per year are diagnosed with CRC (Table 1-2), i.e. 9.0% of incident cancer cases in men and 12.2% in women. CRC was responsible for 78 deaths per year in men from 2007 to 2014, representing 9.1% of all cancer-related deaths, and 74 deaths per year in women (11.2%).

Overall, these 3 regions have similar world-standardized incidence rates, which remain lower than incidence in France as a whole; CRC mortality is also lower.

Liver

In Guadeloupe, Martinique and French Guiana, liver cancer is diagnosed in an average of 30 men and 14 women per year, accounting for 1.4% of incident cancer cases in men, and 1.0% in women. It was

responsible for 38 deaths per year in men from 2007 to 2014, representing 4.4% of cancer-related deaths, and 23 deaths per year in women (3.5% of cancer-related deaths).

Pancreas

In Martinique, Guadeloupe and French Guiana, pancreatic cancer is diagnosed in an average of 44 men and 39 women per year, accounting for 2.0% of incident cancers in men and 2.7% in women. It was responsible for 53 deaths per year between 2007 and 2014, corresponding to 6.2% of cancer deaths in men and 49 deaths per year (7.4% of cancer-related deaths) in women.

Discussion

Overall, Martinique, Guadeloupe and French Guiana present world-standardized incidence rates that vary somewhat between the three regions, but that are in general lower than overall rates for the whole of France; there is also lower mortality related to oesophageal cancer.

The incidence of oesophageal cancer has been declining for several years in men, whereas it is increasing in women.

The majority of oesophageal cancers can be classed into two histological groups. Epidermoid cancers are the most frequent in France in both men and women [12]. Furthermore, the World Health Organization (WHO) has established that X-rays and gamma radiation can contribute to oesophageal cancer. The second histological type is adenocarcinoma; the majority of adenocarcinomas of the oesophagus develop in the context of endobrachyoesophagus (Barrett's oesophagus), following the metaplasia-dysplasia-carcinoma sequence.

In the Caribbean, few data are available on oesophagus cancers incidence and mortality; data available from GLOBOCAN Observatory show that incidence cancer of oesophagus cancer was below 4.0 per 100,000 person-years in men and below 1.0 in women. Latin America and the Caribbean (LAC) region had the lowest incidence rates for this cancer compared to Europe (1.3 per 100,000 in women and 5.8 in men) or Asia (5.3 per 100,000 in women and 12.1 in men). Mortality data in the LAC was 0.91 per 100,000 pers-years in women and 3.7 in men. In our study, our rates are similar to those observed in the LAC [13 14].

We observed incidence and mortality rates of 6.0 and 4.5 per 100,000 pers-years in women for stomach cancer. In men theses rates were respectively 11.0 and 8.8 per 100,000 pers-years. In our study we observed higher incidence and mortality rate, especially in men [14]. The estimated incidence of stomach cancer in France is amongst the lowest of all the estimates provided by the WHO [15]. Incidence of stomach cancer is on the decline over the long term, and this is confirmed by the observations between 2005 and 2012 [16]. Mortality has also been declining persistently since the 1990s[16].

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The risk of stomach cancer is increased in first-degree relatives of a patient with stomach cancer, in individuals who have undergone partial gastrectomy for cancer, or in those who have undergone endoscopic treatment for gastric cancer, as well as in case of precancerous lesions of the stomach and in persons originating from regions with high stomach cancer incidence. Obesity and gastro-oesophageal reflux are risk factors for cardia cancer [17-19]. In Guadeloupe, prevalence of Helicobacter pylori infection, the main risk factor for this type of cancer, was estimated to be 55% in blood donors. Further studies are required to estimate the prevalence in the general population, and among patients with cancer. Environmental risk factors, as well as high intake of salt, and smoked meat and fish in the French West Indies, as in certain Asian countries, could also contribute to the high incidence of stomach cancer. Consumption of fruit and vegetables has a protective effect against stomach cancer.

In the LAC regions, colorectal cancer incidence rates in women was 15.1 per 100,000 pers-years in women and 18.5 in men. Mortality rates were respectively 7.3 per 100,000 pers-years in women and 9.4 in men [14]. In our study, we observed that our incidence and mortality rates were higher compared to Caribbean rates. This could be explained by the development of organized colorectal cancer screening since 2008 in Martinique and Guadeloupe, and in 2009 in French Guiana, that could help to detect more cancer cases. Furthermore, we observed that western lifestyle impacts the incidence and mortality rates in our regions.

Colorectal cancer is also a cancer among those requiring a policy of care, from the stage of organized or individual cancer screening to follow-up as part of the various treatments delivered. Several studies have been carried out in Martinique by the Martinique Cancer Registry, on the evolution of this cancer but also on the factors conditioning survival by age at diagnosis [6 8 9]. Additional studies are underway to analyze regional survival within the French West-Indies and should allow a better understanding of the profile of cancer patients in our region. In France, this cancer benefits from a prevention program which involves health actors from the general practitioner to integration within the framework of the hospital care pathway.

Liver cancer includes primary carcinoma of the liver or hepatocellular carcinoma (HCC, which represents more than 80% of all liver cancers), as well as cancer of the intra-hepatic bile ducts. It is more frequent in France than in Europe as whole or in developed countries [15].

In men, mortality has been declining since 1995, but the opposite trend has been observed in women [16].

There is also a lower mortality from liver cancer in men. Conversely, in women, world-standardized mortality rates show no significant excess- or a lower mortality compared to the rates observed in mainland France.

There are numerous established risk factors for primary liver cancer [20] namely: alcohol consumption, hepatitis B and C viruses, obesity [20], diabetes and tobacco smoking. Bile duct cancer risk factors include liver fluke, biliary tract diseases (primary sclerosing cholangitis and Caroli disease), and exposure to certain chemical compounds such as dichloropropane or dichloromethane[21].

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Geographic variations in incidence observed over the study period could be explained by heterogeneity in the prevalence of chronic diseases linked to alcohol use, hepatitis B and C, or steatosis [22]. Progress in the management of patients with cirrhosis enables carcinogenesis to continue, and also contributes to the increased number of liver cancer cases[22], notably cancer of the intrahepatic bile ducts, whereas other histological types, such as HCC, are on the decline[23]. The distinct geographic distribution of the main risk factors for liver cancer, particularly alcohol consumption, likely explains to a large extent the disparities between Departments in terms of incidence and mortality.

In women, liver cancer incidence and mortality were respectively 4.0 and 3.7 per 100,000 pers-years; in men these rates were respectively 5.9 and 5.6 per 100,000 pers-years in the LAC [14]. In our study our incidence and mortality rates were lower except for French Guiana with an incidence rate of 10.5 per 100,000 pers-years in men.

We observed incidence and mortality rates of 4.0 and 3.8 per 100,000 pers-years in women in the LAC. In men these rates were respectively 5.0 and 4.9 per 100,000 pers-years [14]. In our study, our incidence and mortality rates were higher compared to the LAC.

The incidence of pancreatic cancer is the highest in Eastern Europe, France and Japan[24]. In France, the incidence rate for pancreatic cancer is higher than the average of the 28 EU countries [16]. Pancreatic cancer is one of the 10 most frequent types of cancer, and its incidence increased in both sexes between 2005 and 2012, whereas mortality has remained practically unchanged since the 1980s [16]

Overall, the 3 regions have world-standardized incidence rates that remain lower than those of France as a whole; mortality from pancreatic cancer is also lower. According to Globocan data for LAC (5.0), in men, the incidence rates were similar than the incidence rates in our study except in Martinique (7.0) where it was higher. In women we observed the results between our study (Guadeloupe: 3.0, Martinique: 4.4, French Guiana: 4.8) and LAC (4.0). However, the mortality rates were higher in our study compared to LAC (4.9) in men. In women the mortality rates were similar between LAC (3.8) and data from our study.

Tobacco smoking and obesity are established environmental risk factors for pancreatic cancer. Lowcalorie diets, high alcohol consumption and increased abdominal fat have also been reported to be precipitating factors, while a diet rich in fruit, vegetables and folates, and regular physical exercise are reported to have a protective effect [22].

Conclusion

Collaborative projects to promulgate this expertise will help to improve knowledge of the clinical, demographic, socio-economic or organizational factors that contribute to the heterogeneity of cancer pathologies in the region. Our results are not sufficient to allow trend analysis, but our important to monitor these first results in the future. The next projects of analysis will allow the carrying out of cancer mapping studies as well as the implementation of studies on the risk factors of stomach cancer and the prevalence of Helicobacter Pylori infection in the Antilles. The study of environmental and behavioral

factors is therefore an important issue for a better understanding of the determinants of health and cancer survival.

DECLARATIONS

Authors' contributions

LID, SB, CJ, JVB, JD were major contributors in writing the manuscript, made substantial contributions to conception and design, JP, BBM, JPe, EC, JM, MB, MBR revising it critically for important intellectual content. EC and FRANCIM Network made substantial contributions to conception and design; and revising it critically for important intellectual content. All authors read and approved the final manuscript.

Ethics approval statement

Our study did not involve direct patient contact. Patients were not involved in the design of this study. Regarding patient involvement, cancer cases are identified through multidisciplinary team meetings, through medical records and the registry, according to the Registry procedures (French National authority for the protection of privacy and personal data). Additional approval from ethical committees was not required.

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Conflict interest statement

The authors declare that there are no conflicts of interest.

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Data Sharing

Not applicable.

			Ine	cidence					Mortali			
	Ne	ew cases ¹		WSR ²		SIR ³		Deaths	Fe	WSR ²		SMR ³
Oesophagus									orua			
Guadeloupe	18	[15;21]	6.1	[5.1;7.4]	0.78	[0.65; 0.93]	14	[12;17]	₹ 4.8	[4.0;5.9]	0.89	[0.73; 1.07
Martinique	12	[10;15]	3.8	[3.1;4.8]	0.52	[0.42; 0.63]	10	[7;12]	No 2.8	[2.2;3.8]	0.56	[0.43; 0.7]
French Guiana	4	[2;6]	4.5	[2.7;7.2]	0.56	[0.34 ; 0.88]			N 2.8	[1.6 ; 4.6]	0.52	[0.31; 0.82
Mainland France	3998	[3851 ; 4153]	7.23	[6.95 ; 7.51]			3031	[2993;3070]	5.18	[5.11; 5.25]		
Stomach									vnl			
Guadeloupe	50	[44 ; 55]	14.3	[12.8;16.1]	1.90	[1.70; 2.11]	32	[29;37]	a 9.3	[8.2;10.7]	2.10	[1.85; 2.3]
Martinique	43	[39;48]	12.1	[10.8;13.7]	1.58	[1.42; 1.76]	33	[29;38]	8.9	[7.7;10.3]	2.05	[1.79; 2.3
French Guiana	14	[11;17]	14.6	[11.2; 18.7]	1.90	[1.48; 2.41]	6	[5;8]	ธิ์ 7.3	[5.3 ; 9.8]	1.64	[1.21;2.1
Mainland France	4707	[4561 ; 4859]	7.88	[7.62; 8.14]			2952	[2914;2990]	³ 4.54	[4.48;4.61]		
Colon-Rectum								-	t de la companya de			
Guadeloupe	81	[74;88]	26.2	[24.0 ; 28.6]	0.66	[0.61; 0.72]	34		10.1	[8.9; 11.5]	0.74	[0.66; 0.8
Martinique	90	[83;97]	26.9	[24.8;29.2]	0.70	[0.65; 0.76]	39	[35;44]	<mark>8</mark> 10.8	[9.5; 12.4]	0.80	[0.71;0.9
French Guiana	23	[19;27]	25.0	[20.4;30.3]	0.70	[0.57; 0.84]	5	[3;6]	6.3	[4.4;8.7]	0.46	[0.33;0.6
Mainland France	22828	[22442;23222]	37.8	[37.2;38.5]			8976	[8910;9041]		[12.6; 12.8]		
Liver						N.			ni.c			
Guadeloupe	10	[8;13]	3.6	[2.8;4.7]	0.27	[0.21; 0.33]	15	[12;18]	4.8	[3.9;5.8]	0.50	[0.41; 0.5
Martinique	11	[9;13]	3.3	[2.6;4.3]	0.27	[0.22; 0.33]	18	[15 ; 22]	g 5.4	[4.5;6.7]	0.57	[0.48;0.6
French Guiana	9	[7;12]	10.5	[7.6;14.3]	0.85	[0.62; 1.13]	5		≥ 5.9	[4.2;8.2]	0.66	[0.47; 0.9
Mainland France	6989	[6730;7261]	12.38	[11.91; 12.87]			5739	[5686; 5791]	≝ 9.25	[9.16; 9.34]		
Pancreas									23.			
Guadeloupe	15	[13;19]	4.8	[3.9;6.0]	0.51	[0.42; 0.62]	24	[20;27]	202 7.0	[6.0; 8.2]	0.94	[0.81;1.0
Martinique	24	[20;27]	7.0	[6.0; 8.2]	0.75	[0.65; 0.87]	25	[22 ; 29]	<u>4</u> 6.9	[5.8; 8.2]	0.94	0.80;1.0
French Guiana		[3;7]	5.5	[3.4;8.3]	0.58	[0.37; 0.86]		[3;6]	S 5.8	[4.0; 8.3]	0.69	[0.48; 0.9
Mainland France		[5460; 5705]	9.52	[9.30; 9.74]		-		[4760; 4857]	¥ 7.70	[7.62; 7.79]		·

BMJ Open Table 1. Annual number of new cases and deaths for digestive cancers in men, standardized incidence and mortality rates, standardized incidence and

 Mainland France
 5581 [5460; 5705]
 9.52 [9.50; 9.74]
 4000 [4700, 4057]0 [7.02, 7.05]

 (1) Incidence mainland France: 2007-2016; Guadeloupe : 2008-2014; Martinique : 2007-2014; French Guiana: 2010-2014. (2) World-standardized rates: rates are standardized to the age structure of the world standard population and expressed per 100,000 person-years. (3) Ratios standardized to mainland France.

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	Incidence								Mortality				
	New cases ¹		WSR ²			SIR ³		Deaths	Fel	WSR ²		SMR ³	
Oesophagus									orua				
Guadeloupe	3	[2.0;5.0]	0.9	[0.5;1.5]	0.52	[0.33 ; 0.79]	2	[1;3]	₹ 0.4	[0.2;0.9]	0.45	[0.25; 0.75	
Martinique	2	[2.0;4.0]	0.6	[0.3;1.1]	0.40	[0.24 ; 0.61]	2	[1;4]	80.6	[0.3;1.2]	0.55	[0.32; 0.88	
French Guiana	1	[0.0;3.0]	1.2	[0.4;2.8]	0.82	[0.30; 1.79]	0	[0;1]	№ 0.4	[0.1;1.4]	0.43	[0.09; 1.27	
Mainland France	1072	[1018 ; 1130]	1.49	[1.41; 1.58]			794	[774;813]	0.96	[0.93 ; 0.98]			
Stomach									wnla				
Guadeloupe	32	[28;37]	7.3	[6.3;8.6]	2.29	[2.00; 2.61]	23	[19;26]	ad 4.4	[3.7;5.3]	2.74	[2.35;3.16	
Martinique	34	[30 ; 39]	6.9	[6.0 ; 8.0]	2.31	[2.04 ; 2.60]	22	[19;26]	<u>8</u> 3.7	[3.1;4.7]	2.53	[2.15; 2.94	
French Guiana	8	[5;10]	7.2	[5.0; 10.1]	2.29	[1.62; 3.15]	3	[2;4]	ธิ์ 3.3	[2.1;5.1]	1.79	[1.15; 2.66	
Mainland France	2587	[2508;2670]	3.22	[3.11; 3.34]			1665	[1637; 1694]	1 .71	[1.68; 1.75]			
Colon-Rectum									ttp:				
Guadeloupe	70	[64;76]	17.3	[15.7; 19.2]	0.68	[0.62; 0.75]	29	[26;33]	6 .2	[5.4;7.3]	0.76	[0.66 ; 0.86	
Martinique	85	[79;92]	20.4	[18.8;22.3]	0.80	[0.74 ; 0.86]	41	[36;46]] 7.8	[6.8;9.0]	0.98	[0.87; 1.10	
French Guiana	18	[14;22]	17.4	[13.8;21.7]	0.75	[0.60; 0.92]	4	[3;6]	8 3.8	[2.5;5.5]	0.54	[0.37; 0.77	
Mainland France	19174	[18895 ; 19458]	24.4	[24.0;24.8]			7938	[7877 ; 8000]	g 7.5	[7.4;7.6]			
Liver							~		nj.c				
Guadeloupe	4	[3;6]	1.1	[0.7;1.8]	0.44	[0.30; 0.63]	9	[7;11]	2 .0	[1.5; 2.7]	0.84	[0.65; 1.06	
Martinique	7	[5;9]	1.8	[1.3; 2.7]	0.64	[0.48 ; 0.84]	12	[9 ; 15]	g 2.5	[1.9; 3.4]	1.04	[0.83; 1.29	
French Guiana	3	[2;5]	2.7	[1.5;4.8]	1.21	[0.66 ; 2.02]	2	[1;3]	م 1.8 کچ	[1.0; 3.2]	0.86	[0.47; 1.43	
Mainland France		Not Available					2118	[2087;2151]	≝2.24	[2.20; 2.29]			
Pancreas									23,				
Guadeloupe	12	[9;15]	3.0	[2.3;3.9]	0.43	[0.34; 0.53]	22	[19;26]	2024.8	[4.0;5.7]	0.98	[0.84 ; 1.1.	
Martinique	22	[19;25]	4.4	[3.7;5.4]	0.75	[0.64 ; 0.87]	24	[21;28]	$\frac{12}{4}$ 4.4	[3.6;5.3]	0.97	[0.83; 1.13	
French Guiana	5	[3;7]	4.8	[3.0; 7.4]	0.76	[0.48; 1.13]	3	[2;5]	≥ 3.2	[2.0;5.0]	0.72	[0.47; 1.0	
Mainland France	5278	[5152;5407]	6.46			-	4601	[4554 ; 4648	₩ 4.83	[4.77; 4.89]			

BMJ Open Table 2. Annual number of new cases and deaths for digestive cancers in women, standardized incidence and mortality ates, standardized incidence and

(1) Incidence mainland France: 2007-2016 ; Guadeloupe : 2008-2014 ; Martinique : 2007-2014 ; French Guiana: 2010-2014. (2) World-standardized rates: rates are standardized to the age structure of the world standard population and expressed per 100,000 person-years. (3) Ratios standardized to mainland ected by copyright. France.

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STROBE Statement

	BMJ Open	136/bm
	STROBE Statement Checklist of items that should be included in reports of observational studies	jopen-2020
Item No	Recommendation	20-0471
1	(<i>a</i>) Indicate the study's design with a commonly used term in the title or the abstract (<i>b</i>) Provide in the abstract an informative and balanced summary of what was done and what was found	67 on

	1	(a) Indicate the study's design with a commonly used term in the title or the abstract $\frac{3}{2}$	1
Title and abstract	1	(b) Provide in the abstract an informative and balanced summary of what was done and what was found $\frac{9}{5}$	1
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	2
1 Objectives	3	State specific objectives, including any prespecified hypotheses	2
Methods		State specific objectives, including any prespecified hypotheses	
1 Study design	4	Present key elements of study design early in the paper	3
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up and data collection	3
7 8 9 0		(a) Cohort study—Give the eligibility criteria, and the sources and methods of selection of participants. Bescribe methods of follow-up Case-control study—Give the eligibility criteria, and the sources and methods of case ascertainment and control selection. Give the	3
Participants	6	rationale for the choice of cases and controls <i>Cross-sectional study</i> —Give the eligibility criteria, and the sources and methods of selection of participants	
		(b) Cohort study—For matched studies, give matching criteria and number of exposed and unexposed Case-control study—For matched studies, give matching criteria and the number of controls per case	3
7 Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	3
Data sources/measurement	8*	For each variable of interest, give sources of data and details of methods of assessment (measurement). $\underline{\underline{B}}_{\underline{e}}$ escribe comparability of assessment methods if there is more than one group	3
Bias	9	Describe any efforts to address potential sources of bias	3
Study size	10	Explain how the study size was arrived at	3
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which grouping sewere chosen and why	3
		(a) Describe all statistical methods, including those used to control for confounding	3
7		(b) Describe any methods used to examine subgroups and interactions	3
3		(c) Explain how missing data were addressed	3
⁹ Statistical methods	12	(d) Cohort study—If applicable, explain how loss to follow-up was addressed	3
1		<i>Case-control study</i> —If applicable, explain how matching of cases and controls was addressed <i>Cross-sectional study</i> —If applicable, describe analytical methods taking account of sampling strategy	
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 Section/Topic

Page 23 of 22		BMJ Open	
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Section/Topic	Item No	Recommendation 71	Reported on Page No
Results		S S S S S S S S S S S S S S S S S S S	
Douticinente	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for gligibility, confirmed eligible, included in the study, completing follow-up, and analysed	3-5
Participants	13**	(b) Give reasons for non-participation at each stage	3-5
		(c) Consider use of a flow diagram	3-5
Description data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders	3-5
Descriptive data	14**	(b) Indicate number of participants with missing data for each variable of interest	3-5
		(c) Cohort study—Summarise follow-up time (eg, average and total amount)	3-5
		Cohort study—Report numbers of outcome events or summary measures over time	3-5
Outcome data	15*	<i>Case-control study</i> —Report numbers in each exposure category, or summary measures of exposure	3-5
		Cross-sectional study—Report numbers of outcome events or summary measures	3-5
		(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 👮% confidence interval).	3-5
		Make clear which confounders were adjusted for and why they were included	
Main results	16	(b) Report category boundaries when continuous variables were categorized	3-5
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	3-5
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	Not applicable
Discussion			
Key results	18	Summarise key results with reference to study objectives	5-7
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	5-7
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	5-7
Generalisability	21	Discuss the generalisability (external validity) of the study results	5-7
Other Information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	8
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TITLE PAGE

Burden of gastric and digestive cancers in the French Caribbean: perspectives from populationbased cancer registries of Martinique, Guadeloupe and French Guiana (2007-2014).

Authors' names and affiliations

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Keywords: cancer registry, incidence, Caribbean, mortality. Digestive cancers

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Burden of gastric and digestive cancers in the French Caribbean: perspectives from populationbased cancer registries of Martinique, Guadeloupe and French Guiana (2007-2014).

Abstract

Objectives: Data from population-based cancer registries contribute to improving our knowledge of Digestive cancer trends worldwide. In this study we present cancer incidence and mortality in Guadeloupe, French Guiana and Martinique for the periods 2008-2014, 2010-2014 and 2007-2014 respectively.

Design: Data were extracted from population-based cancer registries. World-standardized incidence and mortality rates were calculated. Main digestive cancers were analysed, including oesophagus, stomach, colorectum, liver and pancreas cancers.

Setting: This study was performed based on data from French Territories in the Caribbean Results:

We observed a lower-incidence compared to mainland France, except for stomach cancer for which the incidence is high, with significant standardized incidence ratios (SIRs) in men and women at 1.90 vs 2.29 for Guadeloupe and French Guiana and 1.58 vs 2.31 for Martinique. We found a lower-mortality, except for stomach cancer for which the mortality remains high, with significant mortality ratios (SMRs) in men and women at 2.10 vs 2.74 for Guadeloupe, 1.64 vs 1.79 for French Guiana and 2.05 vs 2.53 for Martinique. Overall, these 3 regions have similar world-standardized incidence (WSI) and mortality (WSM) rates which remain lower than those in mainland France. We noticed an overall high incidence and high mortality in men compared to women as in France.

Conclusions: There is a high incidence of stomach cancer in French overseas territories. Publication of these data contributed to expanding knowledge on the epidemiology of world cancers with data from the Caribbean zone.

Keywords: cancer registry, incidence, Caribbean, mortality. Digestive cancers **Strengths and limitations of this study**

- The purpose of this project is to present incidence and mortality for digestive cancer in Guadeloupe, French Guiana and Martinique for the 2007-2014 period.
- This study will contribute to expanding knowledge on the epidemiology of world cancers with data from the Caribbean zone.
- Potential limitations include the fact that comorbidities and risk factors are not recorded and thus cannot be taken into account in statistical analyses.

Introduction

A total number of 111 933 new cancer cases were estimated in the Caribbean in 2018 according to GLOBOCAN Database. The most common cancer types were prostate, breast, lung and colorectum cancers; cervical and stomach cancers had higher incidence rates compared to mainland France. Digestive cancer represented 20.6% of these incident cases in both sexes[1].

Main identified risk factors of digestive cancers include socio-economic status, chronic tobacco smoking, and alcoholism. Conversely, a diet rich in fruit and vegetables has been shown to have a protective effect[2]. Other risk factors of oesophageal adenocarcinoma include gastro-oesophageal reflux and obesity [3]. For stomach cancer, *Helicobacter pylori* infection, high intake of salt, exposure to N-nitroso compounds (through diet, tobacco and endogenous synthesis) have been identified as major causes of cancer development[4].

The French West-Indies have a particular socio-demographic profile compared to the Caribbean, with high life expectancy and favourable health indicators. Nevertheless, certain digestive cancers appear in over-incidence such as stomach cancer [4] and underline the need for a study of the evolution of cancers over time from the cancer registries.

The cancer control strategy implemented with the various cancer plans in France [5] has enabled the deployment of significant resources to reduce disparities in the face of cancer. Through the development of health promotion policies, the general population is made aware of the impact of certain risk factors on the development of cancer. Studies are therefore necessary in order to explore the evolution of digestive cancers in our regions [6-9].

Data from population-based cancer registries contribute to improving our knowledge of cancer trends worldwide. The cancer indicators generated are useful for the general population, for researchers, clinicians and local and governmental organisations, and to decision-makers in public health. Pooling of data from the three registries of the French overseas departments will make it possible to identify clinical and epidemiological characteristics of digestive cancers. In this study we present incidence and mortality for digestive cancer in Guadeloupe, French Guiana and Martinique for the 2008-2014, 2010-2014 and 2007-2014 periods respectively.

Methods

Data sources for incidence and mortality

The population-based cancer registries of Martinique, Guadeloupe and French Guiana use patient records which are reviewed actively. The Data quality control procedures are performed according to the French Network of cancer registries FRANCIM and the International Agency for Research cancer (IARC). The control of all available cancer data sources guarantees high quality cancer registration data for international comparisons.

Incidence data

Data were extracted for Guadeloupe, French Guiana and Martinique for the periods 2008-2014, 2010-2014 and 2007-2014 respectively. They are coded according to the International Classification of Diseases for Oncology, Third Edition (ICD-O-3). For mainland France, incidence data (2007-2016 period) were estimated from healthcare and registry data, using a dedicated method described elsewhere [10].

Mortality data

Mortality data cover the period 2007 to 2014. All the data were extracted from the Centre for Epidemiology of the medical causes of death (CepiDC). Data for the year 2012 were not exploitable for Martinique.

Statistical methods

Standardized rates were calculated using the world standard population of the WHO as standard [11]. The standardized incidence ratio (SIR) or standardized mortality ratio (SMR) were calculated in this study, using incidence and mortality rates from mainland France as references. We present the average annual number of cases and deaths observed and the world-standardized incidence and mortality rates by regions, and at national level. The standardized incidence and mortality ratios for these three regions are also presented, with 95% confidence intervals.

Patient and Public Involvement

Our study did not involve direct patient contact. Patients were not involved in the design of this study. Regarding patient involvement, cancer cases are identified through multidisciplinary team meetings, through medical records and the registry, according to the Registry procedures (French National authority for the protection of privacy and personal data). Additional approval from ethical committees was not required.

Results

Main digestive cancers were analysed, including oesophagus, stomach, colorectum, liver and pancreas cancers. In our study, we observed a lower-incidence in all digestive cancer, except for stomach cancer.

Oesophagus

In Martinique, Guadeloupe and French Guiana, oesophageal cancer affects on average 34 men and 6 women per year (Table 1-2), i.e. 1.6% of incident cancer cases in men and 0.4% in women. It was responsible for 26 deaths per year in men between 2007 and 2014 (Table 1), representing 3.0% of cancer deaths, and 4 deaths per year in women (0.6%).

Stomach

In the French West-Indies, 80 individuals were diagnosed with stomach cancer each year respectively in Guadeloupe and Martinique, and 20 in French Guiana. Stomach cancer is more common in these Departments than in mainland France, and represents 4.5% of cancers in men in Martinique, 5.3% in Guadeloupe and 5.7% in French Guiana.

In men, it is the 3rd most common malignancy in Martinique and Guadeloupe, and 4th most common in French Guiana. In women, it is the 3rd most common malignancy in Martinique and 4th most common in Guadeloupe, whereas it is 8th in French Guiana.

Observed incidence is comparable in men in Guadeloupe and French Guiana (respectively 14.3 and 14.6 per 100,000 person-years), and lower in Martinique (12.1). In women, world-standardized incidence rates are 6.9 in Martinique, 7.2 in French Guiana and 7.3 in Guadeloupe. This higher incidence of stomach cancer, with significant standardized incidence ratios (SIRs) at 1.90 for Guadeloupe and French Guiana and 1.58 for Martinique, place these 3 Departments at the top 3 of French regional incidence for this cancer.

In line with incidence data, mortality is also higher than in mainland France. In men, the worldstandardized mortality rate, which reached 9.3 per 100,000 person-years in Guadeloupe, 8.9 in Martinique and 7.3 in French Guiana, is between 1.6 and 2.1 times higher than the corresponding rate in mainland France (4.5) and in most French regions with the exception of Corsica. These differences are also evident for stomach cancer mortality in women, where the world-standard mortality rate of 4.4 per 100,000 person-years in Guadeloupe is more than 2.7 times higher than that of mainland France (1.7). Among the French West-Indies, Guadeloupe has the highest world-standard mortality rates, in both men and women.

Colon-Rectum

In the regions of Guadeloupe, Martinique and French Guiana, on average, 194 men and 173 women per year are diagnosed with CRC (Table 1-2), i.e. 9.0% of incident cancer cases in men and 12.2% in women. CRC was responsible for 78 deaths per year in men from 2007 to 2014, representing 9.1% of all cancer-related deaths, and 74 deaths per year in women (11.2%).

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Liver

In Guadeloupe, Martinique and French Guiana, liver cancer is diagnosed in an average of 30 men and 14 women per year, accounting for 1.4% of incident cancer cases in men, and 1.0% in women. It was responsible for 38 deaths per year in men from 2007 to 2014, representing 4.4% of cancer-related deaths, and 23 deaths per year in women (3.5% of cancer-related deaths).

Pancreas

In Martinique, Guadeloupe and French Guiana, pancreatic cancer is diagnosed in an average of 44 men and 39 women per year, accounting for 2.0% of incident cancers in men and 2.7% in women. It was responsible for 53 deaths per year between 2007 and 2014, corresponding to 6.2% of cancer deaths in men and 49 deaths per year (7.4% of cancer-related deaths) in women.

Discussion

Overall, Martinique, Guadeloupe and French Guiana present world-standardized incidence rates that vary somewhat between the three regions, but that are in general lower than overall rates for the whole of France; there is also lower mortality related to oesophageal cancer.

The incidence of oesophageal cancer has been declining for several years in men, whereas it is increasing in women.

The majority of oesophageal cancers can be classed into two histological groups. Epidermoid cancers are the most frequent in France in both men and women [12]. Furthermore, the World Health Organization (WHO) has established that X-rays and gamma radiation can contribute to oesophageal cancer. The second histological type is adenocarcinoma; the majority of adenocarcinomas of the oesophagus develop in the context of endobrachyoesophagus (Barrett's oesophagus), following the metaplasia-dysplasia-carcinoma sequence.

In the Caribbean, few data are available on oesophagus cancers incidence and mortality; data available from GLOBOCAN Observatory show that incidence cancer of oesophagus cancer was below 4.0 per 100,000 person-years in men and below 1.0 in women. Latin America and the Caribbean (LAC) region had the lowest incidence rates for this cancer compared to Europe (1.3 per 100,000 in women and 5.8 in men) or Asia (5.3 per 100,000 in women and 12.1 in men). Mortality data in the LAC was 0.91 per 100,000 pers-years in women and 3.7 in men. In our study, our rates are similar to those observed in the LAC [13 14].

We observed incidence and mortality rates of 6.0 and 4.5 per 100,000 pers-years in women for stomach cancer. In men theses rates were respectively 11.0 and 8.8 per 100,000 pers-years. In our study we

observed higher incidence and mortality rate compared to mainland France, especially in men [14]. The estimated incidence of stomach cancer in France is amongst the lowest of all the estimates provided by the WHO [15]. Incidence of stomach cancer is on the decline over the long term, and this is confirmed by the observations between 2005 and 2012 in mainland France [16]. Mortality has also been declining persistently since the 1990s [16].

The risk of stomach cancer is increased in first-degree relatives of a patient with stomach cancer, in individuals who have undergone partial gastrectomy for cancer, or in those who have undergone endoscopic treatment for gastric cancer, as well as in case of precancerous lesions of the stomach and in persons originating from regions with high stomach cancer incidence. Obesity and gastro-oesophageal reflux are risk factors for cardia cancer [17-19]. In Guadeloupe, prevalence of Helicobacter pylori infection, the main risk factor for this type of cancer, was estimated to be 55% in blood donors. Further studies are required to estimate the prevalence in the general population, and among patients with cancer. Environmental risk factors, as well as high intake of salt, and smoked meat and fish in the French West Indies, as in certain Asian countries, could also contribute to the high incidence of stomach cancer. Consumption of fruit and vegetables has a protective effect against stomach cancer.

In the LAC regions, colorectal cancer incidence rates in women was 15.1 per 100,000 pers-years in women and 18.5 in men. Mortality rates were respectively 7.3 per 100,000 pers-years in women and 9.4 in men [14]. In our study, we observed that our incidence and mortality rates were higher compared to Caribbean rates. This could be explained by the development of organized colorectal cancer screening since 2008 in Martinique and Guadeloupe, and in 2009 in French Guiana, that could help to detect more cancer cases. Furthermore, we observed that western lifestyle impacts the incidence and mortality rates in our regions.

Colorectal cancer is also a cancer among those requiring a policy of care, from the stage of organized or individual cancer screening to follow-up as part of the various treatments delivered. Several studies have been carried out in Martinique by the Martinique Cancer Registry, on the evolution of this cancer but also on the factors conditioning survival by age at diagnosis [6 8 9]. Additional studies are underway to analyze regional survival within the French West-Indies and should allow a better understanding of the profile of cancer patients in our region. In France, this cancer benefits from a prevention program which involves health actors from the general practitioner to integration within the framework of the hospital care pathway.

Liver cancer includes primary carcinoma of the liver or hepatocellular carcinoma (HCC, which represents more than 80% of all liver cancers), as well as cancer of the intra-hepatic bile ducts. It is more frequent in France than in Europe as whole or in developed countries [15].

In men, mortality has been declining since 1995 in mainland France, but the opposite trend has been observed in women [16].

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There is also a lower mortality from liver cancer in men in our study for the three regions. Conversely, in women, world-standardized mortality rates show no significant excess- or a lower mortality compared to the rates observed in mainland France.

There are numerous established risk factors for primary liver cancer [20] namely: alcohol consumption, hepatitis B and C viruses, obesity [20], diabetes and tobacco smoking. Bile duct cancer risk factors include liver fluke, biliary tract diseases (primary sclerosing cholangitis and Caroli disease), and exposure to certain chemical compounds such as dichloropropane or dichloromethane[21].

Geographic variations in incidence observed over the study period could be explained by heterogeneity in the prevalence of chronic diseases linked to alcohol use, hepatitis B and C, or steatosis [22]. Progress in the management of patients with cirrhosis enables carcinogenesis to continue, and also contributes to the increased number of liver cancer cases[22], notably cancer of the intrahepatic bile ducts, whereas other histological types, such as HCC, are on the decline[23]. The distinct geographic distribution of the main risk factors for liver cancer, particularly alcohol consumption, may explain the disparities between Departments in terms of incidence and mortality.

In women, liver cancer incidence and mortality were respectively 4.0 and 3.7 per 100,000 pers-years; in men these rates were respectively 5.9 and 5.6 per 100,000 pers-years in the LAC [14]. In our study our incidence and mortality rates were lower except for French Guiana with an incidence rate of 10.5 per 100,000 pers-years in men.

We observed incidence and mortality rates of 4.0 and 3.8 per 100,000 pers-years in women in the LAC. In men these rates were respectively 5.0 and 4.9 per 100,000 pers-years [14]. In our study, our incidence and mortality rates were higher compared to the LAC.

The incidence of pancreatic cancer is the highest in Eastern Europe, France and Japan[24]. In France, the incidence rate for pancreatic cancer is higher than the average of the 28 EU countries [16]. Pancreatic cancer is one of the 10 most frequent types of cancer, and its incidence increased in both sexes between 2005 and 2012, whereas mortality has remained practically unchanged since the 1980s [16]

Overall, the 3 regions have world-standardized incidence rates that remain lower than those of France as a whole; mortality from pancreatic cancer is also lower. According to Globocan data for LAC (5.0), in men, the incidence rates were similar than the incidence rates in our study except in Martinique (7.0) where it was higher. In women we observed the results between our study (Guadeloupe: 3.0, Martinique: 4.4, French Guiana: 4.8) and LAC (4.0). However, the mortality rates were higher in our study compared to LAC (4.9) in men. In women the mortality rates were similar between LAC (3.8) and data from our study.

Tobacco smoking and obesity are established environmental risk factors for pancreatic cancer. Lowcalorie diets, high alcohol consumption and increased abdominal fat have also been reported to be precipitating factors, while a diet rich in fruit, vegetables and folates, and regular physical exercise are reported to have a protective effect [22].

Conclusion

Collaborative projects to promulgate this expertise will help to improve knowledge of the clinical, demographic, socio-economic or organizational factors that contribute to the heterogeneity of cancer burden in the region. Our results are not sufficient to allow trend analysis, but our important to monitor these first results in the future. The next projects of analysis will allow the carrying out of cancer mapping studies as well as the implementation of studies on the risk factors of stomach cancer and the prevalence of Helicobacter Pylori infection in the Antilles. The study of environmental and behavioral factors is therefore an important issue for a better understanding of the determinants of health and cancer survival.

DECLARATIONS

Authors' contributions

LID, SB, CJ, JVB, JD were major contributors in writing the manuscript, made substantial contributions to conception and design, JP, BBM, JPe, EC, JM, MB, MBR revising it critically for important intellectual content. EC and FRANCIM Network made substantial contributions to conception and design; and revising it critically for important intellectual content. All authors read and approved the final manuscript.

Ethics approval statement

Our study did not involve direct patient contact. Patients were not involved in the design of this study. Regarding patient involvement, cancer cases are identified through multidisciplinary team meetings, through medical records and the registry, according to the Registry procedures (French National authority for the protection of privacy and personal data). Additional approval from ethical committees was not required.

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2 3	Conflict interest statement
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5 6	The authors declare that there are no conflicts of interest.
6 7	Source of funding
8	Santé publique France, Institut national du cancer. COP N°2019-071
9 10	Data Sharing
11	Not applicable.
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	Incidence									Mortality		
	Ne	ew cases ¹		WSR ²		SIR ³		Deaths	Г e	WSR ²		SMR ³
Oesophagus									orua			
Guadeloupe	18	[15;21]	6.1	[5.1;7.4]	0.78	[0.65; 0.93]	14		4 .8	[4.0;5.9]	0.89	[0.73; 1.07
Martinique	12	[10;15]	3.8	[3.1;4.8]	0.52	[0.42; 0.63]	10	[7;12]	8 2.8	[2.2;3.8]	0.56	[0.43; 0.71
French Guiana	4	[2;6]	4.5	[2.7; 7.2]	0.56	[0.34; 0.88]	2	[1;4]	^N 2.8	[1.6;4.6]	0.52	[0.31; 0.82
Mainland France	3998	[3851 ; 4153]	7.23	[6.95 ; 7.51]			3031	[2993; 3070]	5.18	[5.11; 5.25]		
Stomach									wnl			
Guadeloupe	50	[44 ; 55]	14.3	[12.8;16.1]	1.90	[1.70; 2.11]	32	[29;37]	a 9.3	[8.2; 10.7]	2.10	[1.85; 2.37
Martinique	43	[39;48]	12.1	[10.8;13.7]	1.58	[1.42; 1.76]	33	[29;38]	8.9	[7.7;10.3]	2.05	[1.79 ; 2.33
French Guiana	14	[11;17]	14.6	[11.2; 18.7]	1.90	[1.48; 2.41]	6	[5;8]	ธิ์ 7.3	[5.3 ; 9.8]	1.64	[1.21; 2.17
Mainland France	4707	[4561 ; 4859]	7.88	[7.62; 8.14]			2952	[2914;2990]	³ 4.54	[4.48; 4.61]		
Colon-Rectum								-	tto:			
Guadeloupe	81	[74;88]	26.2	[24.0 ; 28.6]	0.66	[0.61; 0.72]	34	[30;39]	10.1	[8.9; 11.5]	0.74	[0.66 ; 0.84
Martinique	90	[83;97]	26.9	[24.8;29.2]	0.70	[0.65; 0.76]	39	[35;44]	i 10.8	[9.5; 12.4]	0.80	[0.71; 0.90
French Guiana	23	[19;27]	25.0	[20.4;30.3]	0.70	[0.57; 0.84]	5		6.3	[4.4 ; 8.7]	0.46	0.33; 0.64
Mainland France		[22442;23222]	37.8	[37.2;38.5]			8976	[8910;9041]	12.7	[12.6; 12.8]		2
Liver								•	Di. C			
Guadeloupe	10	[8;13]	3.6	[2.8;4.7]	0.27	[0.21; 0.33]	15	[12;18]	4.8	[3.9;5.8]	0.50	[0.41; 0.59
Martinique	11	[9;13]	3.3	[2.6; 4.3]	0.27	[0.22; 0.33]	18	[15;22]	g 5.4	[4.5;6.7]	0.57	0.48;0.68
French Guiana	9	[7;12]	10.5	[7.6; 14.3]	0.85	[0.62; 1.13]		[3;7]	≥ 5.9	[4.2;8.2]	0.66	0.47; 0.91
Mainland France	6989	[6730 ; 7261]	12.38	[11.91 ; 12.87]			5739	[5686; 5791]	9.25 9 .25	[9.16; 9.34]		
Pancreas									23.			
Guadeloupe	15	[13;19]	4.8	[3.9;6.0]	0.51	[0.42; 0.62]	24	[20;27]	20 7.0	[6.0; 8.2]	0.94	[0.81 ; 1.08
Martinique	24	[20;27]	7.0	[6.0; 8.2]	0.75	[0.65 ; 0.87]	25	[22;29]	4 6.9	[5.8; 8.2]	0.94	0.80; 1.09
French Guiana		[3;7]	5.5	[3.4; 8.3]	0.58	[0.37; 0.86]			5.8	[4.0; 8.3]	0.69	0.48; 0.96
Mainland France		[5460; 5705]	9.52	[9.30; 9.74]				[4760; 4857]	¥ 7.70			

BMJ Open Table 1. Annual number of new cases and deaths for digestive cancers in men, standardized incidence and mortality rates, standardized incidence and

(1) Incidence mainland France: 2007-2016 ; Guadeloupe : 2008-2014 ; Martinique : 2007-2014 ; French Guiana: 2010-2014. (2) World-standardized rates: rates are standardized to the age structure of the world standard population and expressed per 100,000 person-years. (3) Ratios standardized to mainland ected by copyright. France.

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			Inc	idence					Mortali			
	N	lew cases ¹		WSR ²		SIR ³		Deaths	Fel	WSR ²		SMR ³
Oesophagus									orua			
Guadeloupe	3	[2.0;5.0]	0.9	[0.5;1.5]	0.52	[0.33 ; 0.79]	2	[1;3]	ຊື 0.4	[0.2;0.9]	0.45	[0.25]
Martinique	2	[2.0;4.0]	0.6	[0.3;1.1]	0.40	[0.24 ; 0.61]	2	[1;4]	≥0.6	[0.3;1.2]	0.55	[0.32 ;
French Guiana	1	[0.0;3.0]	1.2	[0.4;2.8]	0.82	[0.30; 1.79]	0	[0;1]	^N 0.4	[0.1;1.4]	0.43	[0.09]
Mainland France	1072	[1018 ; 1130]	1.49	[1.41; 1.58]			794	[774;813]	0.96	[0.93 ; 0.98]		
Stomach									n			
Guadeloupe	32	[28;37]	7.3	[6.3;8.6]	2.29	[2.00; 2.61]	23	[19;26]	ad 4.4	[3.7;5.3]	2.74	[2.35 ;
Martinique	34	[30 ; 39]	6.9	[6.0 ; 8.0]	2.31	[2.04 ; 2.60]	22	[19;26]	<u>8</u> 3.7	[3.1;4.7]	2.53	[2.15 ;
French Guiana	8	[5;10]	7.2	[5.0; 10.1]	2.29	[1.62;3.15]	3	[2;4]	ธิ์ 3.3	[2.1;5.1]	1.79	[1.15;
Mainland France	2587	[2508 ; 2670]	3.22	[3.11; 3.34]			1665	[1637;1694]	<u> </u>] <u>-</u> 1.71	[1.68; 1.75]		
Colon-Rectum									-te			
Guadeloupe	70	[64;76]	17.3	[15.7 ; 19.2]	0.68	[0.62; 0.75]	29	[26;33]	6 .2	[5.4;7.3]	0.76	[0.66;
Martinique	85	[79;92]	20.4	[18.8;22.3]	0.80	[0.74 ; 0.86]	41	[36;46]] 7.8	[6.8;9.0]	0.98	[0.87;
French Guiana	18	[14;22]	17.4	[13.8;21.7]	0.75	[0.60; 0.92]	4	[3;6]	8 3.8	[2.5;5.5]	0.54	[0.37;
Mainland France	19174	[18895 ; 19458]	24.4	[24.0;24.8]			7938	[7877;8000]] <mark>5</mark> 7.5	[7.4;7.6]		
Liver									nj.c			
Guadeloupe	4	[3;6]	1.1	[0.7;1.8]	0.44	[0.30; 0.63]	9	[7;11]	2 .0	[1.5;2.7]	0.84	[0.65;
Martinique	7	[5;9]	1.8	[1.3;2.7]	0.64	[0.48 ; 0.84]	12	[9 ; 15]	g 2.5	[1.9;3.4]	1.04	[0.83;
French Guiana	3	[2;5]	2.7	[1.5;4.8]	1.21	[0.66 ; 2.02]	2	[1;3]	₽1.8	[1.0; 3.2]	0.86	[0.47;
Mainland France		Not Available					2118	[2087 ; 2151]] ⊒2.24	[2.20; 2.29]		
Pancreas									23,			
Guadeloupe	12	[9;15]	3.0	[2.3;3.9]	0.43	[0.34 ; 0.53]	22	[19;26]	824.8	[4.0;5.7]	0.98	[0.84 ;
Martinique	22	[19;25]	4.4	[3.7;5.4]	0.75	[0.64 ; 0.87]	24	[21 ; 28]	44	[3.6;5.3]	0.97	[0.83 ;
French Guiana	5	[3;7]	4.8	[3.0;7.4]	0.76	[0.48;1.13]	3	[2;5]	by 3.2	[2.0;5.0]	0.72	[0.47 ;
Mainland France	5278	[5152;5407]	6.46	[6.29; 6.63]			4601	[4554 ; 4648] آه 4.83	[4.77 ; 4.89]		

BMJ Open Table 2. Annual number of new cases and deaths for digestive cancers in women, standardized incidence and mortality ates, standardized incidence and

(1) Incidence mainland France: 2007-2016 ; Guadeloupe : 2008-2014 ; Martinique : 2007-2014 ; French Guiana: 2010-2014. (2) World-standardized rates: rates are standardized to the age structure of the world standard population and expressed per 100,000 person-years. (3) Ratios standardized to mainland ected by copyright. France.

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		BMJ Open STROBE Statement Checklist of items that should be included in reports of observational studies
Section/Topic	Item No	Recommendation 47
Title and abstract	1	(<i>a</i>) Indicate the study's design with a commonly used term in the title or the abstract
	1	(b) Provide in the abstract an informative and balanced summary of what was done and what was found $\frac{9}{8}$
Introduction		
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported
1 Objectives	3	State specific objectives, including any prespecified hypotheses
² Methods		
4 Study design	4	Present key elements of study design early in the paper Q
5 6 Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up and data collection $\frac{1}{2}$
7 8 9 0 1 Participants 2 3 4 5	6	 (a) Cohort study—Give the eligibility criteria, and the sources and methods of selection of participants. Bescribe methods of follow-up Case-control study—Give the eligibility criteria, and the sources and methods of case ascertainment and control selection. Give the rationale for the choice of cases and controls Cross-sectional study—Give the eligibility criteria, and the sources and methods of selection of participants. (b) Cohort study—For matched studies, give matching criteria and number of exposed and unexposed Case-control study—For matched studies, give matching criteria and the number of controls per case

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25		Case-control study—For matched studies, give matching criteria and the number of controls per case	
26 27 Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if	2
27 Variables 28	8*	applicable S	
29		For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of	3
³⁰ Data sources/measurement	8.	assessment methods if there is more than one group	
31 32 Bias	9	Describe any efforts to address potential sources of bias	3
33 Study size	10	Explain how the study size was arrived at	3
4 Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which grouping were chosen and why	3
35 36		(a) Describe all statistical methods, including those used to control for confounding	3
57		(b) Describe any methods used to examine subgroups and interactions	3
8		(c) Explain how missing data were addressed	3
⁹ Statistical methods	12	(d) Cohort study—If applicable, explain how loss to follow-up was addressed	3
.0 .1 .2		<i>Case-control study</i> —If applicable, explain how matching of cases and controls was addressed	
		Cross-sectional study—If applicable, describe analytical methods taking account of sampling strategy	
43		(e) Describe any sensitivity analyses	Not
14			1

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11 Objectives

14 Study design

16 Setting

ge 19 of 18		BMJ Open	
		jop en-20	applicable
Section/Topic	Item No	Recommendation 47167	Reported on Page No
Results		S S S S S S S S S S S S S S S S S S S	
Destisingute	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for gligibility, confirmed eligible, included in the study, completing follow-up, and analysed	3-5
Participants	13**	(b) Give reasons for non-participation at each stage	3-5
		(c) Consider use of a flow diagram	3-5
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders	3-5
Descriptive data	14**	(b) Indicate number of participants with missing data for each variable of interest	3-5
		(c) Cohort study—Summarise follow-up time (eg, average and total amount)	3-5
		Cohort study—Report numbers of outcome events or summary measures over time	3-5
Outcome data	15*	Case-control study—Report numbers in each exposure category, or summary measures of exposure	3-5
		Cross-sectional study—Report numbers of outcome events or summary measures	3-5
		(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 👮% confidence interval).	3-5
Main namlta	16	Make clear which confounders were adjusted for and why they were included	
Main results	16	(b) Report category boundaries when continuous variables were categorized	3-5
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	3-5
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	Not applicable
Discussion			
Key results	18	Summarise key results with reference to study objectives	5-7
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	5-7
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	5-7
Generalisability	21	Discuss the generalisability (external validity) of the study results	5-7
Other Information		ec	
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	8
	y for cases	and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross \vec{s} ctional studies.	
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