ORIGINAL ARTICLE

Global patterns and trends in colorectal cancer incidence and mortality

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ABSTRACT

Objective The global burden of colorectal cancer (CRC) is expected to increase by 60% to more than 2.2 million new cases and 1.1 million deaths by 2030. In this study, we aim to describe the recent CRC incidence and mortality patterns and trends linking the findings to the prospects of reducing the burden through cancer prevention and care.

Design Estimates of sex-specific CRC incidence and mortality rates in 2012 were extracted from the GLOBOCAN database. Temporal patterns were assessed for 37 countries using data from Cancer Incidence in Five Continents (CI5) volumes I-X and the WHO mortality database. Trends were assessed via the annual percentage change using joinpoint regression and discussed in relation to human development levels. **Results** CRC incidence and mortality rates vary up to 10-fold worldwide, with distinct gradients across human

development levels, pointing towards widening disparities and an increasing burden in countries in transition. Generally, CRC incidence and mortality rates are still rising rapidly in many low-income and middleincome countries: stabilising or decreasing trends tend to be seen in highly developed countries where rates remain among the highest in the world.

Conclusions Patterns and trends in CRC incidence and mortality correlate with present human development levels and their incremental changes might reflect the adoption of more western lifestyles. Targeted resourcedependent interventions, including primary prevention in low-income, supplemented with early detection in highincome settings, are needed to reduce the number of patients with CRC in future decades.

INTRODUCTION

Colorectal cancer (CRC) is the third most commonly diagnosed malignancy and the fourth leading cause of cancer death in the world, accounting for about 1.4 million new cases and almost 700 000 deaths in 2012.1 The distribution of CRC burden varies widely, with more than two-thirds of all cases and about 60% of all deaths occurring in countries with a high or very high human development index (HDI).1 CRC is considered one of the clearest markers of the cancer transition, replacing infection-related cancers in countries undergoing rapid societal and economic changes together with other cancers predominantly linked to western lifestyles, which are already frequently found in high-income countries.^{2–4}

Significance of this study

What is already known on this subject?

- Colorectal cancer (CRC) is the third most commonly diagnosed malignancy and the fourth leading cause of cancer-related deaths in the world, and its burden is expected to increase by 60% to more than 2.2 million new cases and 1.1 million cancer deaths by 2030.
- CRC is considered one of the clearest markers of the cancer transition, whereby countries undergoing rapid societal and economic changes show rapid increases in cancers already more frequent in high-income countries.
- Understanding the current patterns of CRC and its evolution from an international perspective is imperative in order to direct future prospects of reducing the burden through cancer prevention and cancer care.

What are the new findings?

- CRC incidence and mortality rates vary widely worldwide, with distinct gradients across human development levels and trends point towards widening disparities and an increasing burden in countries in transition.
- We identified three patterns of CRC incidence and mortality trends and ascertained that CRC incidence and mortality rates are still rising rapidly in many low-income and middle-income countries, linked to ongoing societal and economic development; stabilising or decreasing trends tend to be seen only in highly developed countries where rates remain among the highest in the world.

How might it impact on clinical practice in the foreseeable future?

- Without targeted resource-dependent actions based on this evidence, the number of patients with CRC will continue to increase in future
- ▶ Improvements in treatment options and accessibility are vital, particularly in low-income and middle-income countries that face an increasing burden of CRC.
- Prioritisation of primary prevention and early detection are necessary, alongside their integration into existing healthcare plans.



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Rapid increases in both CRC incidence and mortality are now observed in many medium-to-high HDI countries particularly in Eastern Europe, Asia and South America.² In contrast, CRC incidence and mortality rates have been stabilising or declining in a number of the highest indexed HDI countries: the USA, Australia, New Zealand and several Western European countries.² The reasons for the recent declining trends in incidence in these countries are ill-defined and likely numerous but may partially reflect increased early detection and prevention through polypectomy (at least in the USA). Together with the factors that have brought about declines in incidence, improvements in perioperative care, as well as chemotherapy and radiotherapy, will have contributed to the uniformly decreasing trends in CRC mortality in many highincome settings.⁵

Given the temporal profiles and demographic projections, the global burden of CRC is expected to increase by 60% to more than 2.2 million new cases and 1.1 million cancer deaths by 2030. Understanding the current patterns of CRC and its

evolution from an international perspective is therefore imperative, and in this study, we describe the geographical variations in CRC incidence and mortality in 184 countries and time trends in 37 countries, linking the findings to the future prospects of reducing the burden through cancer prevention and care.

METHODS

Incidence and mortality estimates of malignant neoplasms of the colon and rectum (ICD-10 C18-21) by country for 184 countries in 2012 were extracted from the GLOBOCAN database. Data on HDI for the same year were obtained from the United Nations Development Programme.

To assess time trends in CRC incidence and mortality, we used data from two different sources with the requirement of at least 15 consecutive years of data, and the availability of both incidence and mortality data for each country included. For CRC incidence, data series from high-quality regional and national population-based cancer registries were extracted from

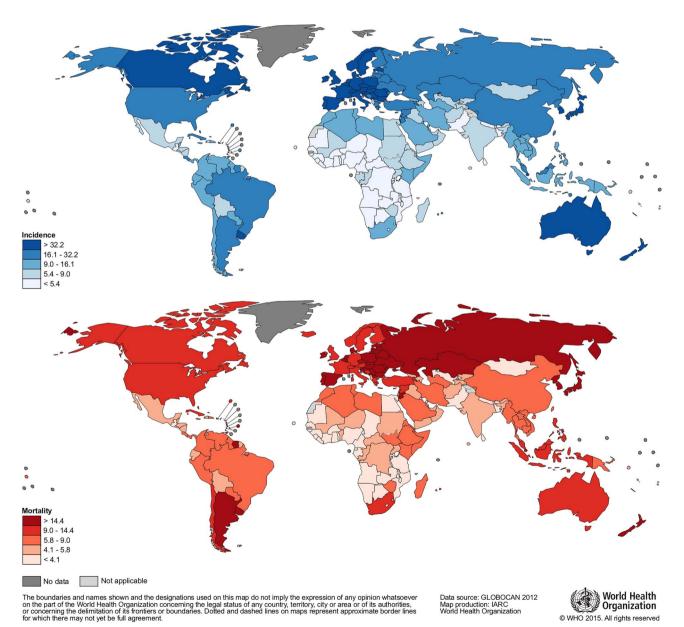


Figure 1 Worldwide colorectal cancer incidence and mortality rates (age adjusted according to the world standard population, per 100 000) in males in 2012 (GLOBOCAN 2012¹).

the *Cancer Incidence in Five Continents* (CI5) series Volumes I–X, ⁸ complemented with publically available data from European countries, ⁹ ¹⁰ ¹¹ Australia, ¹² New Zealand, ¹³ and the USA ¹⁴ for more recent years. Of the 37 countries studied, national incidence data were available for 25 countries. For the remaining countries, data from regional registries were pooled to obtain a proxy of the national incidence (see online supplementary annex table S1). National mortality data series were extracted from the WHO mortality database, with the minimal inclusion criteria set at the WHO-defined medium data quality level, ensuring a reasonable degree of population coverage, completeness and accuracy. ¹⁵ Rates were age-standardised (ASR) to the world standard population. ¹⁶

To analyse incidence and mortality trends, we used joinpoint regression, ¹⁷ which involves fitting a series of joined straight lines to ASR trends. A logarithmic transformation of the rates, calculation of SEs using the binomial approximation, and a maximum number of three joinpoints were specified as options in the analysis. To estimate the magnitude and direction of recent trends, we calculated the average annual percentage change (AAPC) and the corresponding 95% CI for the last available 10 years. The AAPC is a geometrically weighted average of the different annual percentage changes from the joinpoint trend analysis, for which weights are equal to the length of each segment during the specified time interval.

RESULTS

Incidence and mortality patterns of CRC in 2012

In 2012, the estimated incidence rates in males varied from <5 (per 100 000) in several African countries to over 40 in certain countries in Europe, Northern America and Oceania (figure 1). The highest rates in males were observed in Slovakia (61.6), Hungary (58.9) and the Republic of Korea (58.7), while the lowest were seen in sub-Saharan Africa, in The Gambia and Mozambique (both 1.5 per 100 000). Geographical patterns were very similar between the sexes, although rates in females tended to be lower (25% less) than their male counterparts (data not shown, see ref. ¹). Gradients in incidence were

observed with increasing levels of HDI: incidence rates in countries with a very high HDI were six times greater than countries with a low HDI (figure 2A). Geographical patterns of CRC mortality rates generally followed those of incidence, although the highest rates observed tended to be in countries with high rather than very high HDI in Central and Eastern Europe and Latin America (figure 1). As with incidence, mortality showed a distinct gradient across HDI levels (figure 2B), while comparisons of incidence-to-mortality revealed higher case fatalities among countries indexed with lower levels of HDI.

Trends in incidence and mortality from CRC

Based on temporal characteristics of incidence and mortality (in males), three different groups of countries were identified (table 1): those with increasing or stable incidence and mortality (group 1, n=14 countries), those with increasing incidence and decreasing mortality (group 2, n=14 countries) and those with decreasing incidence and mortality (group 3, n=9 countries). The results are presented according to these three categories.

Group 1: increasing incidence and mortality

Increases in both incidence and mortality over the most recent 10-year period were seen in this group, comprising several Eastern European countries, and also in populations in Latin America and Asia (see figures 3A and 4A/B; online supplementary annex tables S2 and S3). In males, the largest increases in incidence were seen in Brazil (AAPC 7.2, 95% CI -7.5 to 24.2), Costa Rica (3.6, 95% CI 3.1 to 4.2) and Bulgaria (3.6, 95% CI 3.1 to 4.2), while mortality rates rose most rapidly in the Philippines (5.7, 95% CI 4.7 to 6.7) and Belarus (3.4, 95% CI 2.5 to 4.3). Incidence uniformly rose in all countries within this group, while mortality rates appeared to level off in Bulgaria, Russia, Croatia, Spain, Latvia and Estonia. Trends in females were similar to those in males, although both incidence and mortality were generally lower except in Latin American countries (Brazil, Costa Rica and Colombia), where rates in males and females were quite similar.

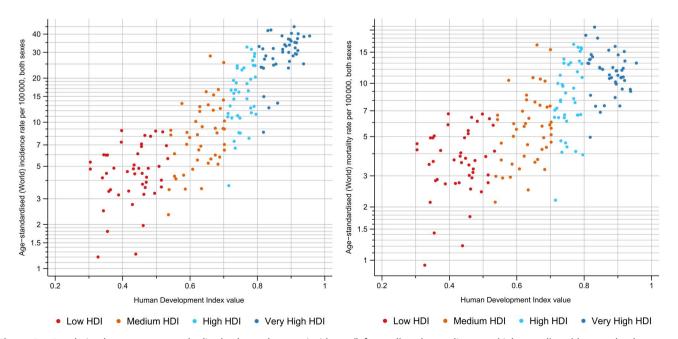


Figure 2 Correlation between age-standardised colorectal cancer incidence (left panel) and mortality rates (right panel) and human development index (HDI) in both sexes combined (GLOBOCAN 2012¹).

 Table 1
 Trends in colorectal cancer incidence and mortality: three groups of temporal pattern

Group 1: incidence ↑ mortality ↑	Philippines*, China*, Colombia†, Bulgaria†, Costa Rica†, Brazil†, Russia†, Belarus†, Estonia, Lithuania, Croatia, Spain, Latvia, Poland
Group 2: incidence ↑ mortality ↓	Canada, Denmark, Switzerland, Ireland, Sweden, Singapore, Finland, Norway, Slovakia, UK, Netherlands Italy, Malta, Slovenia
Group 3: incidence ↓ mortality ↓	US (White), US (Black), Austria, New Zealand, Czech Republic, Iceland, France, Japan, Australia, Israel

^{*}Medium human development index (HDI), refers to $0.534 < \text{HDI} \le 0.710$.

Group 2: increasing incidence and decreasing mortality

In this group, incidence rates continued to increase while mortality rates decline, an observation seen in several European countries, and also in Canada and Singapore (see figures 3B and 4A/B; online supplementary annex tables S2 and S3). The rate of increase in incidence in males was marked in Southern European countries including Slovenia (2.7, 95% CI 0.5 to 5.0), Malta (1.7, 95% CI 0.5 to 2.9) and Italy (1.7, 95% CI 1.5 to 1.9), while stable or slight rises in trends were observed in

Northern European countries and in Canada. Although mortality rates decreased most remarkably in Ireland (-2.1, 95% CI -2.9 to -1.3) and Denmark (-1.9, 95% CI -2.3 to -1.6), the corresponding incidence rates remained constant during the most recent 10-year period. As declining mortality rates were also observed in the UK (-2.0, 95% CI -2.2 to -1.8), these were, however, paralleled by a significant average increase of 1.3% per year in incidence. Trends were similar in both sexes.

Group 3: decreasing incidence and mortality

Decreases in both CRC incidence and mortality in this group were restricted to the highest HDI countries such as Australia, Iceland, New Zealand and Japan (see figures 3C and 4A/B; online supplementary annex tables S2 and S3). While the magnitude of declining incidence in males were marked among US Whites (AAPC -3.0, 95% CI -3.3 to -2.7) and Blacks (-2.2, 95% CI -3.3 to -1.0), mortality declines were greatest (-4.6% per annum) in the Czech Republic. Again, patterns were similar for males and females.

DISCUSSION

This paper provides a comprehensive overview of the patterns and trends in the overall CRC incidence and mortality worldwide. CRC incidence and mortality rates varied widely, with

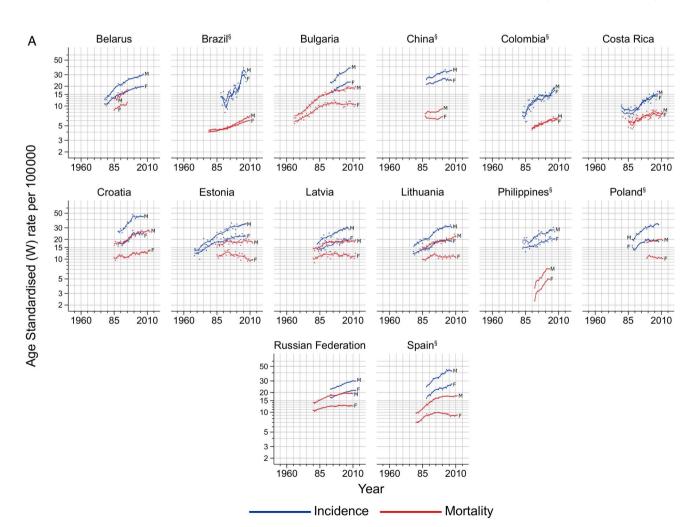
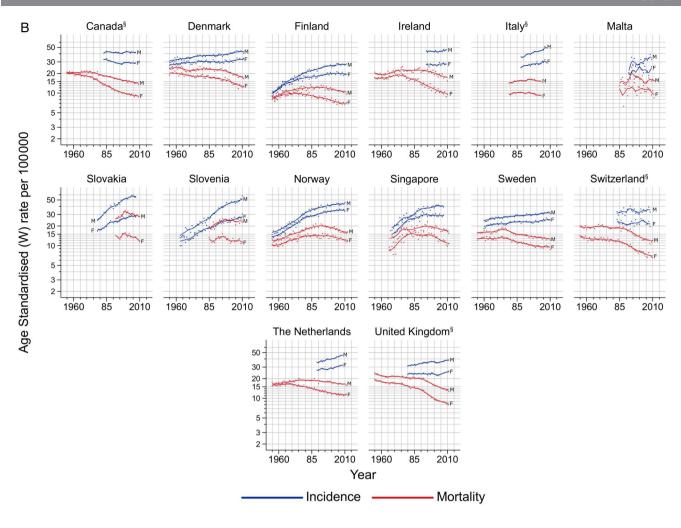


Figure 3 (A) Trends in colorectal cancer incidence and mortality in males (M) and females (F) by country (group 1: increasing or stable incidence and mortality). §Regional data. (B) Trends in colorectal cancer incidence and mortality in males (M) and females (F) by country (group 2: increasing incidence and decreasing mortality). §Regional data. (C) Trends in colorectal cancer incidence and mortality in males (M) and females (F) by country (group 3: decreasing incidence and mortality). §Regional data.

Very high HDI (all remaining countries) refers to HDI > 0.796.



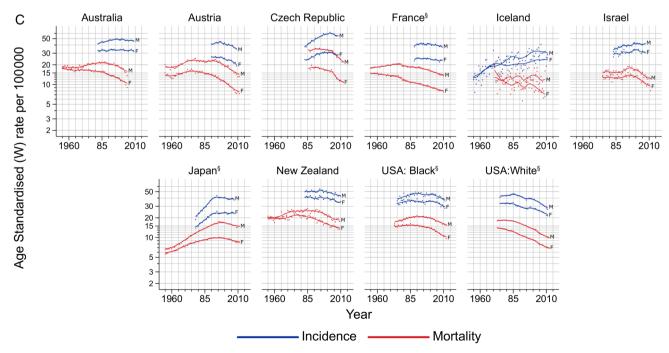


Figure 3 Continued

distinct gradients across HDI levels. We identified three patterns of CRC incidence and mortality trends: (1) increases in both incidence and mortality in the most recent decade, as observed in rapidly transitioning countries (eg, medium and high HDI countries including those in the Baltics, as well as Russia, China and Brazil), (2) increases in incidence, with concomitant decreases in mortality, as seen in very high HDI countries including Canada, UK, Denmark and Singapore and (3) decreases in both incidence and mortality, as observed in a number of the highest HDI-indexed countries including the USA, Japan and France.

The mortality declines observed in latter two groups affect both sexes and have been previously reported in North America (the USA and Canada^{5 18}), Oceania (Australia and New Zealand), most European countries (other than Croatia, Romania, Latvia, Estonia and Russia) and Asia (Japan). These can partly be linked to improving survival through the adoption of best practices in cancer treatment and management for CRC. 19 20 Removal of polyps and early detection efforts, 21-24 including the adoption of colonoscopy, flexible sigmoidoscopy, CT colonography, faecal immunochemistry testing and faecal occult blood testing, may be responsible for the rest of the observed trends. The introduction of screening tests may initially increase CRC incidence rates due to the detection of undiagnosed disease but has been shown to reduce incidence longer term due to the removal of precancerous polyps during colonoscopy.²⁴ This may be particularly pertinent in explaining the uniformly decreasing mortality trends in the USA, Israel and Japan, countries where organised screening and early detection programmes have been established since the 1990s. 25 However, the extent to which screening interventions are responsible for the recent reduction in incidence rates in these countries, relative to a reduction in risk via a changing prevalence and distribution of the key risk factors, is difficult to clarify at present. Other high-income countries have introduced organised screening practices very recently; it is however unlikely that screening has materially influenced recent incidence trends in these

The presence of birth cohort effects implies the importance of changing risk in successive generations in contributing to the recent plateau or declines in incidence observed in certain very high HDI countries without long-standing organised screening programmes, most notably Australia, New Zealand and several European countries.²⁶ Certainly, changes in the prevalence of lifestyle-related factors linked to the extent of 'westernisation' are likely to be, in part, responsible for the global variation in CRC incidence as well as the observed increasing incidence trends in countries in transition in groups (1) and (2). These modifiable risk factors include alcohol consumption, ^{27–29} poor diet (low consumption of fruits and vegetables, and high consumption of red/processed meats), 30-36 obesity, 37 38 physical inactivity³⁹ 40 and smoking. 41–43 Despite higher relative risks, family history of CRC44 and IBD45 accounts for only a small proportion of the observed variation of CRC burden globally given their lower prevalence. Established protective factors that could partly explain stabilising of incidence rates in high-income countries include the regular use of aspirin, 46 47 the use of oestrogens after menopause⁴⁸ and possibly vitamin D intake.⁴⁹

Dietary patterns and the overall composition of diet have shifted dramatically over the past half-century, with distinct differences within world regions and individual countries. In Japan, for example, cereal consumption decreased sharply and vegetable consumption remained almost stable since the 1950s until 1990.⁵⁰ In contrast, meat and fat intake increased sharply

from the mid-1950s until the early-1970s to mid-1970s. The slow increase until the last decade has been met with marked concomitant increases in both colon cancer incidence and mortality during the years 1990–2000. The rapid transition in income and economic growth in low-income and middle-income countries has shifted dietary patterns towards an increased intake of fat, sugar and animal-source foods. Changes in the food environment including access to cheaper 'junk' food were also paralleled by reductions in physical activity and increases in sedentary behaviour, fuelled by both increases in overweight and obesity and changes in the built environment. The second sec

Increases in mortality have been reported in several countries in Latin America, the Caribbean and Asia, 54 55 and these may reflect limited health infrastructure and poorer access to early detection and treatment.⁵⁶ Survival from CRC depends heavily on the stage at diagnosis,^{57–59} and the unfavourable distribution of advanced cancers in low-income and middle-income countries may explain the higher M:I ratios as well as increases in mortality in these countries. For colon cancer, typically the tumour and corresponding lymph vessels are removed during surgery and adjuvant chemotherapy is administered to patients at high risk of relapse. 60 As for rectal cancer, complete removal of the mesorectum is the standard surgical procedure that has been shown to increase survival and substantially decrease the risk of recurrence.⁶⁰ ⁶¹ In addition, typically a combination of (neo)adjuvant chemotherapy and radiotherapy is administered, whereby the recommended regimen depends heavily on the tumour type and stage at diagnosis. Yet, in low-income settings such as sub-Saharan Africa, surgery is often the only available treatment option and adjunctive therapy often not available.⁶² Among all patients with cancer receiving radiotherapy in lowincome and middle-income countries only 1.3% and 3.1% received radiotherapy for cancers of the colon and rectum, respectively, while the 'optimum' proportion should have been 14% and 61%, respectively.⁵⁸ Furthermore, delays in diagnosis, referral and treatment and also cultural beliefs and financial constraints, for example, in rural areas of Latin America, may explain part of the higher mortality in this region. 62 63

This study has a number of strengths and limitations. We have aimed to provide a comprehensive analysis of geographical variations of both CRC incidence and mortality in 2012 by sex and the corresponding trends in both indicators in 37 countries. Attention should be drawn to lack of availability of recorded (registry) incidence and mortality data (particularly in low-income and middle-income countries) in deriving national incidence and mortality estimates in 184 countries as part of our GLOBOCAN compilation; only one-third and one-fifth of the world's countries presently report high-quality incidence and mortality data, respectively. For the trends analyses, we used CI5 data of high comparability, completeness and validity to assess trends in incidence. In using national mortality, we used only data with at least WHO-defined medium levels of completeness and coverage.

The intention was to provide a global snapshot of the scale and profile of CRC today, using high-quality data wherever possible. This study serves as a pointer to show how the disease burden is likely to develop in low-income countries in the longer term and highlights the pressing need for cancer control action to halt the rising mortality rates in many low-income and middle-income countries. While the scope of this study necessitates a general approach, the main weakness stems from a lack of granularity in the analyses. We have not included here a separate assessment of colon and rectal cancer,

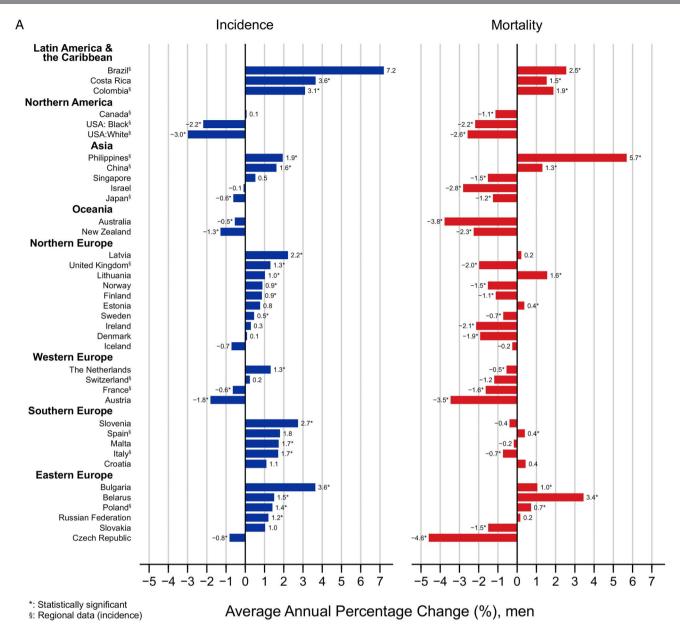


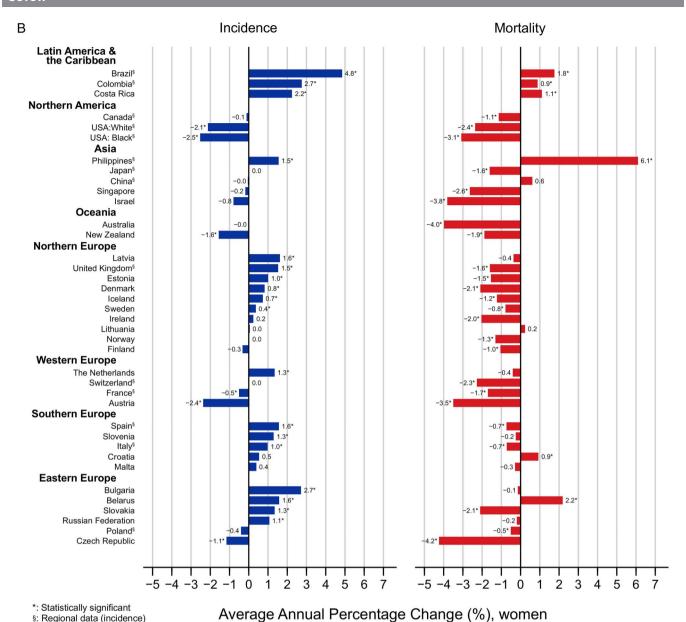
Figure 4 (A) Average annual percentage change (AAPC) of colorectal cancer incidence and mortality in the most recent period (10 years), males. (B) AAPC of colorectal cancer incidence and mortality in the most recent period (10 years), females.

nor did we examine variations in distal (left-sided) cancers of the rectosigmoid junction and proximal (right-sided) cancers of the descending and sigmoid colon, despite many studies reporting a rising proportion of the latter tumours. The inclusion of birth cohort analyses would also have led to a more robust assessment of the recent direction of trends enabling the generation of hypotheses linked to the changing prevalence of causative factors in successive cohorts and the impact of early-stage factors and early-in-life experiences, and their further study will increase our understanding of the aetiology of this cancer.

Diverse global CRC patterns and trends point towards widening disparities and an increasing burden in countries in transition. Generally, CRC incidence and mortality rates correlate with the adoption of a western lifestyle; while they are still rising rapidly in many low-income and middle-income countries linked to ongoing societal and economic development, in highly developed countries, rates are stabilising or decreasing.

Decreases in incidence (also seen in recent birth cohorts) in high HDI and high-risk countries are likely driven by changes in life-style and dietary patterns over the past decades, which might translate into further future rate declines as these cohorts age. Early detection and screening might have led to short-term increases in incidence, but such interventions will eventually contribute to mortality reductions through the increased detection of early-stage tumours.

The fact that CRC has replaced infection-related cancers as the second most common cancer in several middle-income countries (particularly among women) highlights the major challenge of CRC control in countries undergoing significant socioeconomic transition, and the importance of continued efforts to monitor trends in CRC incidence, mortality and survival worldwide. Without targeted resource-dependent actions based on this evidence, the number of patients with CRC will continue to increase in future decades beyond those already projected as a result of population ageing and population growth.



§: Regional data (incidence)

Figure 4 Continued

Improvements in treatment options and accessibility, including tertiary care, are vital in low-income and middle-income countries that face an increasing burden of CRC. In light of the limited capacity for health service provision and associated costs, however, there remains an overwhelming need to prioritise and integrate primary prevention and early detection measures into existing healthcare plans.⁶⁴

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GUT

10-fold difference worldwide in new cases of, and deaths from, bowel cancer

Rising rates linked to economic development; adoption of Western lifestyle may be to blame

There's a 10-fold difference worldwide in the numbers of new cases of bowel cancer and deaths from the disease, finds research published online in the journal *Gut.*

International bowel cancer patterns and trends seem to be linked to economic development, the analysis shows, suggesting that the adoption of a Western lifestyle may have a role.

Bowel cancer is the third most commonly diagnosed cancer and the fourth leading cause of cancer death in the world. In 2012 there were an estimated 1.4 million new cases and almost 700,000 associated deaths worldwide.

By 2030, the numbers of new cases are expected to surge to 2.2 million with an associated death toll of 1.1 million.

In a bid to provide a global snapshot of the international distribution of bowel cancer, and the prospects for curbing predicted rises in rates, the researchers extracted data from the GLOBOCAN database on the numbers of new cases and deaths from the disease in 2012 for 184 countries.

They also looked at time trends in 37 countries, using data from 10 volumes of *Cancer Incidence in Five Continents* and the World Health Organisation mortality database.

The estimated rates of new cases in 2012 varied from less than 5 per 100,000 of the population in several African countries to over 40 per 100,000 in certain countries in Europe, Northern America, and Oceania.

Rates in women tended to be around 25% lower than those of men, among whom the highest rates were in Slovakia (61.6), Hungary (58.9), and Korea (58.7). The lowest rates (1.5/100,000) were in sub-Saharan Africa, The Gambia, and Mozambique.

Using information from the United Nations Development Programme, bowel cancer rates were analysed across levels of economic development, referred to as the HDI, for 2012 in all 184 countries.

The higher the HDI, the higher was the incidence of bowel cancer. In countries with a very high HDI the number of new cases was, on average, six times higher than in countries with a very low HDI.

Patterns of deaths from the disease tended to follow those of incidence, although the highest rates tended to be in countries with high, rather than very high, HDI, in central and eastern Europe and in Latin America.

The time trends analysis revealed three distinct groups of countries: those with rising or stable incidence and death rates (group 1, 14 countries); those with rising incidence and falling death rates (group 2, 14 countries); and those with falling incidence and death rates (group 3, 9 countries).

Group 1 included The Philippines, China, Columbia, Bulgaria, Costa Rica, Brazil, Russia, Belarus, Estonia, Lithuania, Croatia, Spain, Latvia, and Poland. Russia, China, Brazil and the Baltics have undergone rapid economic development over the past decade, the researchers note.

Group 2 included Canada, Denmark, Switzerland, Ireland, Sweden, Singapore, Finland, Norway, Slovakia, the UK, Netherlands, Italy, Malta, and Slovenia.

Group 3 included the USA, Austria, New Zealand, Czech Republic, Iceland, France, Japan, Australia and Israel. But in these countries rates remain among some of the highest in the world.

The researchers suggest that the fall in deaths in groups 2 and 3 is partly linked to improved detection and treatment of the disease.

But the findings point to a much greater disease toll in low and middle income countries in the years to come, particularly for emerging economies, they warn.

The study was observational, so it can increase our understanding of possible links between global patterns, economic development and bowel cancer, but it does not prove cause and effect because other factors may play a role.

"The fact that [bowel cancer] has replaced infection related cancers as the second most common cancer in several middle income countries (particularly among women) highlights the major challenge of [bowel cancer] control in countries undergoing significant socioeconomic transition," they write.

"Without targeted resource dependent actions based on this evidence, the number of patients with bowel cancer will continue to increase in future decades beyond those already projected as a result of population ageing and population growth," they say.

Table 1. Regional registries (proportion of national population covered)

Country	Registries (coverage)
Brasil	Goiânia (~1%)
Canada	All provinces but Quebec(~75%)
China	Hong-Kong and Shanghai(~1%)
Colombia	Cali (~5%)
France	Bas-Rhin, Calvados, Doubs, Isere, Haut-Rhin, Herault, Somme and Tarn (~10%)
Italy	City of Turin and Modena, Parma, Romagna, Ragusa, Varese provinces (~7%)
Japan	Miyagi, Nagasaki and Osaka (~8%)
Philippines	Manila (~7%)
Poland	Cracow city, Kielce and Lower Silesia (~13%)
Spain	Granada, Murcia, Navarra and Tarragona (~8%)
Switzerland	Geneva and St. Gall-Appenzell (~13%)
Thailand	Chiang Mai (~3%)
United Kingdom	England and Wales (~93%)
USA: black	SEER 9 (~9%)
USA: White	SEER 9 (~9%)

Table 2. Number of new colorectal cancer cases, average population year, age-standardized incidence rates per 100,000 (ASR) for the period 2003-2007 and average annual percentage change (AAPC) for the most recent 10-year period.

Country			Males				1	Females		AAPC
	Cases	Population	ASR	AAPC	CI (95%)	Cases	Population	ASR	AAPC CI (95%)	period
Northern Europe										
Denmark	10346	2682224		0.1	(-1.4)	9673	2737817	31.9	(0.4)	- 1
Estonia	1638	620195		8.0	(-0.4)	1911	726350	22.1	(0.5)	- 1
Finland	6 237	2568387		*6.0	(0.7)	8809	2680076	19.9	(-1.0)	- 1
Iceland	374	150602		-0.7	(-2.3)	305	148087	24.4	(0.4	- 1
Ireland	6 138	2067940	43.9	0.3	(-0.1)	4641	2078440	27.9	$0.2 (-0.2 \ ; 0.7)$	2000 - 2009
Latvia	2424	1059911		2.2*	(1.9)	2876	1240634	19.9	(1.1	- 1
Lithuania	3672	1592521		1.0^{*}	(0.3)	3770	1822239	19.4	(-1.2)	- 1
Norway	8664	2297763		*6.0	(0.7	8 871	2332246	35.2	(-1.1	- 1
Sweden	14409	4483716		0.5*	(0.4)	13634	4558280	24.8	(0.3)	- 1
$\operatorname{United\ Kingdom}^{\S}$	82075	24753540		1.3*	(1.0; 1.6)	68574	25700440	23.4	1.5*(1.0;2.0)	- 1
Western Europe										
Austria	13712	3993909	39.2	-1.8*	(-2.3;-1.3)	11404	4222465	22.8	-2.4* $(-3.5;-1.2)$	2000 - 2009
${ m France}^{ m S}$	10164	2939121	39.5	+9.0-	(-0.9;-0.3)	8092	3112434	23.3	-0.5*(-0.7;-0.3)	2000 - 2009
Netherlands	29249	8068312	43.4	1.3*	(1.2;1.5)	26203	8242627	30.9	1.3^* (1.1; 1.6)	1999 - 2008
${ m Switzerland}^{\S}$	1418	465736	35.5	0.2	(-0.2;0.7)	1230	490401	22.8	$0.0 (-0.4 \ ; 0.4)$	1999 - 2008
Southern Europe										
Croatia	8 140	2138032	44.5	1.1	(-0.4; 2.6)	6311	2301798	24.6	$0.5 (-1.5 \; ; \; 2.6)$	
$Italy^{\$}$	10802	2080603	47.0	1.7*	(1.5;1.9)	8764	2201980	29.9	1.0* (0.7; 1.3)	1998 - 2007
Malta	520	200308	32.2	1.7*	(0.5; 2.9)	408	203473	21.4	$0.4 (-1.1 \ ; 1.9)$	
Slovenia	3690	983120	46.4	2.7*	(0.5;5.0)	2877	1021443	25.6	1.3* (0.8; 1.8)	
${ m Spain}^{\$}$	6355	1753690	43.4	1.8	(-0.2;3.9)	4486	1744913	25.6	1.6* (1.1; 2.0)	1998 - 2007
Eastern Europe										
Belarus	8 771	4572634	29.0	1.5*	(1.1	9939	5209029	19.7	(1.3)	1998 - 2007
Bulgaria	11965	3756129	34.8	3.6*	(3.1	9552	3984553	22.3	(2.4	1999 - 2008
Czech Republic	23131	4998500	57.5	-0.8*	(-1.6 ; -0.0)	16368	5247900	29.2	-1.1* $(-1.8; -0.5)$	1999 - 2008
$\mathrm{Poland}^{\$}$	5 7 5 7	2371098	33.0	1.4^{*}	(0.6)	4918	2567624	19.2	(-1.1]	1999 - 2008
Russian Federation	117502	66120721	28.0	1.2*	(0.6)	146755	76420546	20.3	(0.8)	2003 - 2012
Slovakia	9621	2615651	26.7	1.0	(-1.0)	7 004	2772730	28.2	(1.1	1998 - 2007

^{§:} Regional data.*: Statistically significant.CI: Confidence interval.

Table 2. Number of new colorectal cancer cases, average population year, age-standardized incidence rates per 100,000 (ASR) for the period 2003-2007 and average annual percentage change (AAPC) for the most recent 10-year period.

Country			Males]	Females		AAPC
Course	Cases	Cases Population	ASR	AAPC	CI (95%)	Cases	Population	ASR	AAPC CI (95%)	period
Latin America										
& the Caribbean										
${ m Brazil}^{\S}$	648	568640	31.5	7.2	(-7.5 ; 24.2)	727	625374	27.5	4.8* (3.1;6.6)	1998 - 2007
Colombia [§]	716	961993	16.7	3.1*	(2.3;3.9)	924	1079001	16.0	2.7* (1.9; 3.6)	1998 - 2007
Costa Rica	1363	2164476	15.1	3.6*	3.6* (3.1;4.2)	1408	2098568	14.2	2.2* (1.7; 2.8)	1998 - 2007
Northern America										
$\operatorname{Canada}^{\S}$	39934	12205078	41.9	0.1	(-0.9;1.0)	33895	12415665	28.7	-0.1 (-0.7; 0.5)	1998 - 2007
$USA: Black^{\S}$	3328	1648251	42.6	-2.2*	(-3.3 ; -1.0)	3729	1819566	34.3	-2.5*(-3.7;-1.3)	2002 - 2011
$USA: White^{\S}$	26745	10352048	33.8	-3.0*	(-3.3 ; -2.7)	26123	10492238	25.5	-2.1*(-2.4;-1.9)	2002 - 2011
Asia										
$China^{\S}$	18657	6380478	33.8	1.6*	(1.3;1.9)	15732	6617649	24.9	$-0.0 (-0.7 \ ; \ 0.7)$	1998 - 2007
Israel	8239	3311974	41.3	-0.1	(-0.9;0.7)	7831	3379817	31.1	$-0.8 (-2.2 \ ; 0.6)$	1998 - 2007
Japan [§]	25285	6180246	38.6	*9.0-	(-0.9;-0.4)	19916	6605365	23.6	$0.0 (-0.2 \ ; 0.3)$	1998 - 2007
${ m Philippines}^{\S}$	2054	2874967	27.8	1.9*	(1.5;2.4)	1957	2981213	20.6	1.5*(1.1;2.0)	1998 - 2007
Singapore	3890	1722740	40.3	0.5	(-0.1;1.1)	3363	1748380	28.8	-0.2 (-0.6 ; 0.2)	1998 - 2007
Oceania										
Australia	37272	10150633	46.9	-0.5*	-0.5*(-0.8;-0.3)	31030	10286951	33.2	-0.0 (-0.1; 0.1)	2000 - 2009
New Zealand	7 019	2 021 788	46.2	-1.3*	(-1.7;-0.9)	6 793	2096262	37.2	-1.6* (-2.3; -0.8)	2001 - 2010

^{§:} Regional data.*: Statistically significant.

CI: Confidence interval.

Table 3. Number of new colorectal cancer deaths, average population year, age-standardized mortality rates per 100,000 (ASR) for the period 2003-2007 and average annual percentage change (AAPC) for the most recent 10-year period.

Country			Males			Н	Females		AAPC
	Cases	Population	ASR	AAPC CI (95%)	Cases	Population	ASR	AAPC CI (95%)	period
Northern Europe									
Denmark	4923	2684306	18.8		4923	2737208	13.3	(-2.6;	2002 - 2011
Estonia	921	612322	18.6	(0.0;	1055	713382	10.9	(-2.2;	- 1
Finland	2601	2568989	10.9	(-1.5	2627	2680610	7.2	(-1.2;	- 1
Iceland	144	149336	12.2	(-1.0	129	147763	8.6	(-1.8;	- 1
Ireland	2705	2071871	18.3	-2.1*(-2.9;-1.3)	2024	2082346	10.5	-2.0*(-2.2;-1.9)	2001 - 2010
Latvia	1563	1021000	18.8	(-0.1	1898	1206500	11.9	(-0.8;	- 1
Lithuania	2249	1527484	20.1	(1.3;	2288	1757986	10.8	(-0.4;	- 1
Norway	3 770	2297947	17.2	(-1.8;	4010	2334636	13.0	(-1.8;	- 1
Sweden	6521	4482285	13.0	(-0.8;	6397	4558121	9.6	(-1.0;	- 1
United Kingdom	37516	26194859	14.4	(-2.2;	32868	27220829	9.0	(-2.1;	- 1
Western Europe									
Austria	6240	3999390	16.8		5690	4233197	9.5	(-3.8;	- 1
France	43516	29670254	14.8	• •	38759	31735999	8.5	-1.7*(-2.0;-1.4)	2001 - 2010
Netherlands	11702	8061155	16.8		11367	8234079	11.5	(-1.0;	- 1
Switzerland	4398	3631831	12.4	(-3.7;	3836	3786002	7.3	(-2.5;	- 1
Southern Europe									
Croatia	4861	2115027	25.9		3607	2274546	12.3	(0.7;	- 1
$Italy^a$	44868	27744064	15.3	(-1.3;	39575	29541641	9.3		1994 - 2003
Malta	245	206374	14.5		233	208612	11.6	(-1.0	- 1
Slovenia	1911	980781	23.7	(-1.2;	1597	1022316	12.1	(-1.6)	- 1
Spain	37460	21344204	17.5	$0.4^* (0.1; 0.7)$	27543	22019029	9.1	-0.7*(-1.1;-0.4)	2002 - 2011
Eastern Europe									
$\mathrm{Belarus}^a$	4480	4797846	16.3	3.4^* (2.5; 4.3)	5114	5440365	10.6	(1.5;	1986 - 1995
Bulgaria	6756	3741165	18.6	1.0* (0.8; 1.3)	5125	3943000	10.6	-0.1 (-0.5 ; 0.2)	2003 - 2012
Czech Republic	12387	4997613	30.1	-4.6*(-5.2;-4.0)	9136	5253402	14.5	(-6.6)	2003 - 2012
Poland	25360	18484797	19.1	0.7* (0.4; 1.0)	22201	19728307	10.6	(-0.9)	2002 - 2011
Russian Federation	82069	66 847 699	19.5	0.2 (-0.1; 0.5)	100467	254	12.6	$-0.2 (-0.6 \ ; 0.2)$	2001 - 2010
Slovakia	cZ1 c	2 017 867	29.8	-1.5* (-2.3; -0.7)	3 087	2776210	13.0	-2.1** (-2.8; -1.4)	2001 - 2010

^{*:} Statistically significant. CI: Confidence interval.

 $[^]a\mathrm{ASR}$ period: Italy 1999-2003, Belarus 1991-1995.

Table 3. Number of new colorectal cancer deaths, average population year, age-standardized mortality rates per 100,000 (ASR) for the period 2003-2007 and average annual percentage change (AAPC) for the most recent 10-year period.

Country			Males			Ħ	Females		AAPC
f Torrison	Cases	Population	ASR	AAPC CI (95%)	Cases	Population	ASR	AAPC CI (95%)	period
Latin America									
& the Caribbean									
Brazil	23928	91762179	6.2	2.5*(2.3;2.7)	27165	94245207	5.7	$1.8^* (1.6; 1.9)$	2001 - 2010
Colombia	4824	21276343	6.3	1.9*(1.5;2.2)	5 783	21908349	0.9	0.9*(0.0;1.7)	2000 - 2009
Costa Rica	922	2194874	8.2	1.5* (0.9; 2.1)	803	2123791	7.6	$1.1^* (0.6; 1.6)$	2002 - 2011
Northern America									
Canada	19584	15992723	14.6	-1.1*(-1.2;-1.0)	17041	16278395	9.2	-1.1*(-1.4;-0.8)	2000 - 2009
USA: Black	16762	18742161	17.9	-2.2* $(-2.5;-1.9)$	17519	20565336	12.3	-3.1* $(-3.4; -2.7)$	2003 - 2012
USA: White	115531	117684798	11.6	-2.6*(-3.2;-2.0)	113412	120477199	8.0	-2.4^* $(-3.0; -1.7)$	2003 - 2012
Asia									
$China^a$	25380	58051537	8.7	$1.3^* (0.9; 1.7)$	21447	55689238	9.9	$0.6 (-0.0 \ ; 1.2)$	1991 - 2000
Israel	3152	3266959	14.4	-2.8*(-3.3;-2.3)	3175	3354970	11.1	-3.8* (-4.8; -2.8)	2002 - 2011
Japan	110929	61936089	15.8	-1.2*(-1.4;-1.1)	93221	64995935	9.2	-1.6^* $(-1.9; -1.3)$	2002 - 2011
$\operatorname{Philippines}^a$	6 708	39925045	8.9	5.7* (4.7;6.7)	5488	39380144	4.7		1994 - 2003
Singapore	1716	1752280	17.5	-1.5*(-2.3;-0.8)	1496	1780100	12.0	-2.6*(-3.6:-1.7)	2002 - 2011
Oceania									
$\mathrm{Australia}^a$	12222	9788335	15.9	-3.8* (-4.7; -2.8)	10403	9938237	10.5	-4.0* $(-5.4;-2.5)$	1995 - 2004
New Zealand	2 947	2023962	18.5	-2.3* (-2.7; -1.8)	3 010	2106369	14.7	-1.9* (-2.2; -1.6)	2000 - 2009

*: Statistically significant. CI: Confidence interval.

 $[^]a\mathrm{ASR}$ period: China 1996-2000, Philippines 1999-2003, Australia 2000-2004.