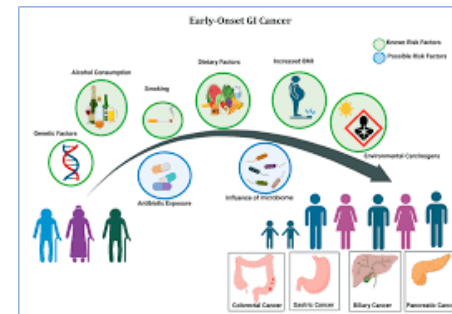
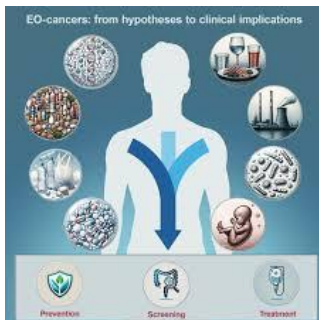


Early-Onset Gastrointestinal Cancers (EOGIC)



الجمعية السورية لأمراض الهضم
Syrian Society of Gastroenterology

المؤتمر العلمي السادس والثلاثين لأمراض الهضم
12 - 14 تشرين الثاني 2025



Early-onset (EO) gastrointestinal (GI) cancer

- Early-onset gastrointestinal (GI) cancer is typically defined as GI cancer diagnosed in individuals younger than 50 years



Trends in
Colorectal
new

RESEARCH

ORIGINAL ARTICLE

Saad El
<https://c>

Fanny ER Vu
Mário Dinis-
Michal F Ka
Marcis Leja

ORIGINAL ART
Increasing
adults in
Global patterns and trends in colorectal cancer
incidence in young adults

Rebecca L Siegel ¹, Lindsey A Torre,¹ Isabelle Soerjomataram,² Richard B Hayes,³
Freddie Bray,² Thomas K Weber,^{4,5} Ahmedin Jemal¹

Global Increasing Incidence of Young-Onset
Colorectal Cancer Across 5 Continents: A
Joinpoint Regression Analysis of 1,922,167 Cases

N. Lui¹, Kelvin K.F. Tsoi^{2,3}, Jason M.W. Ho³, C.M. Lo³, Felix C.H. Chan³,
dorp-Vogelaar,²
sje,⁸ Laura Esteban,⁹
ndřej Májek,^{14,15}

young

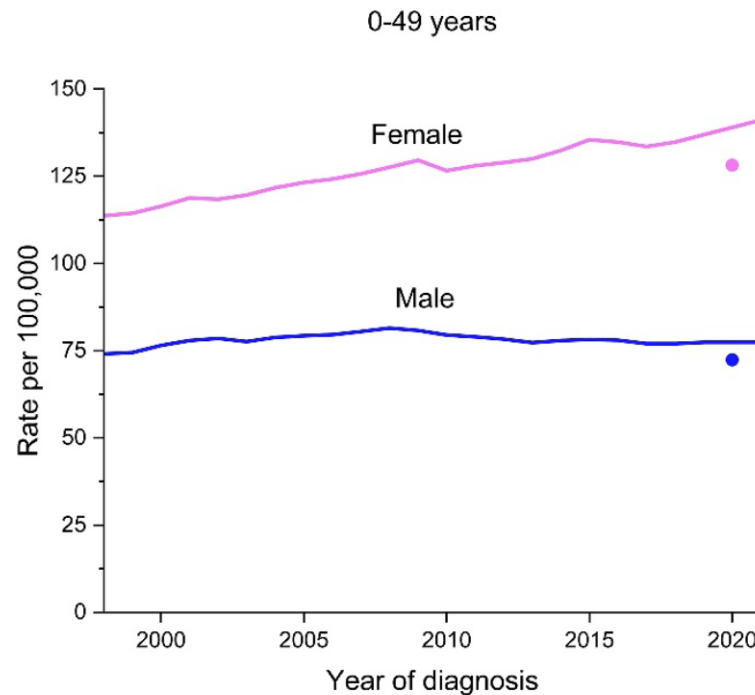
Cancer
Epidemiology,
Biomarkers
& Prevention



Young-onset
stema

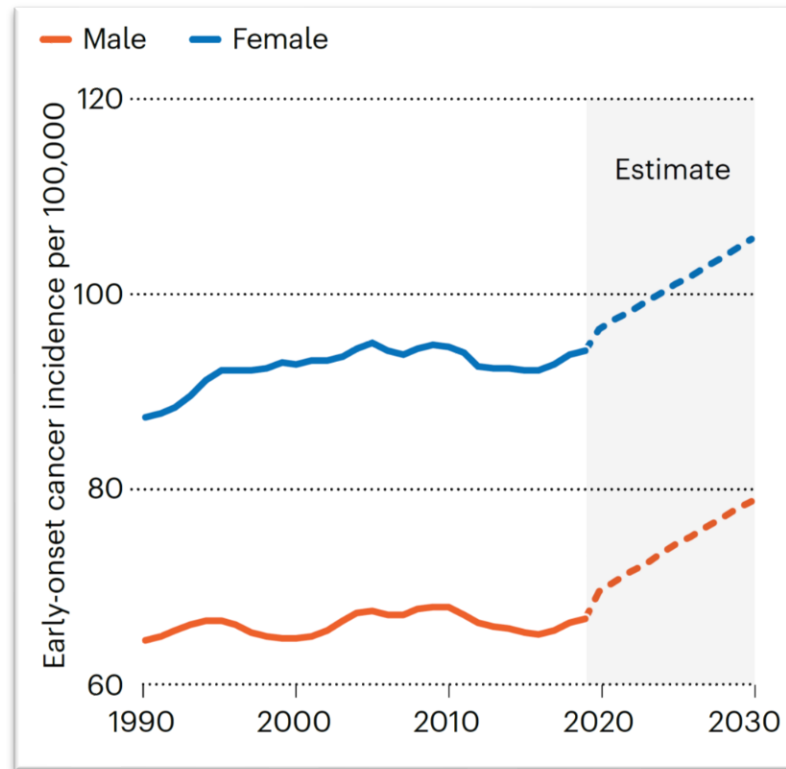
er

Cancer statistics, 2025



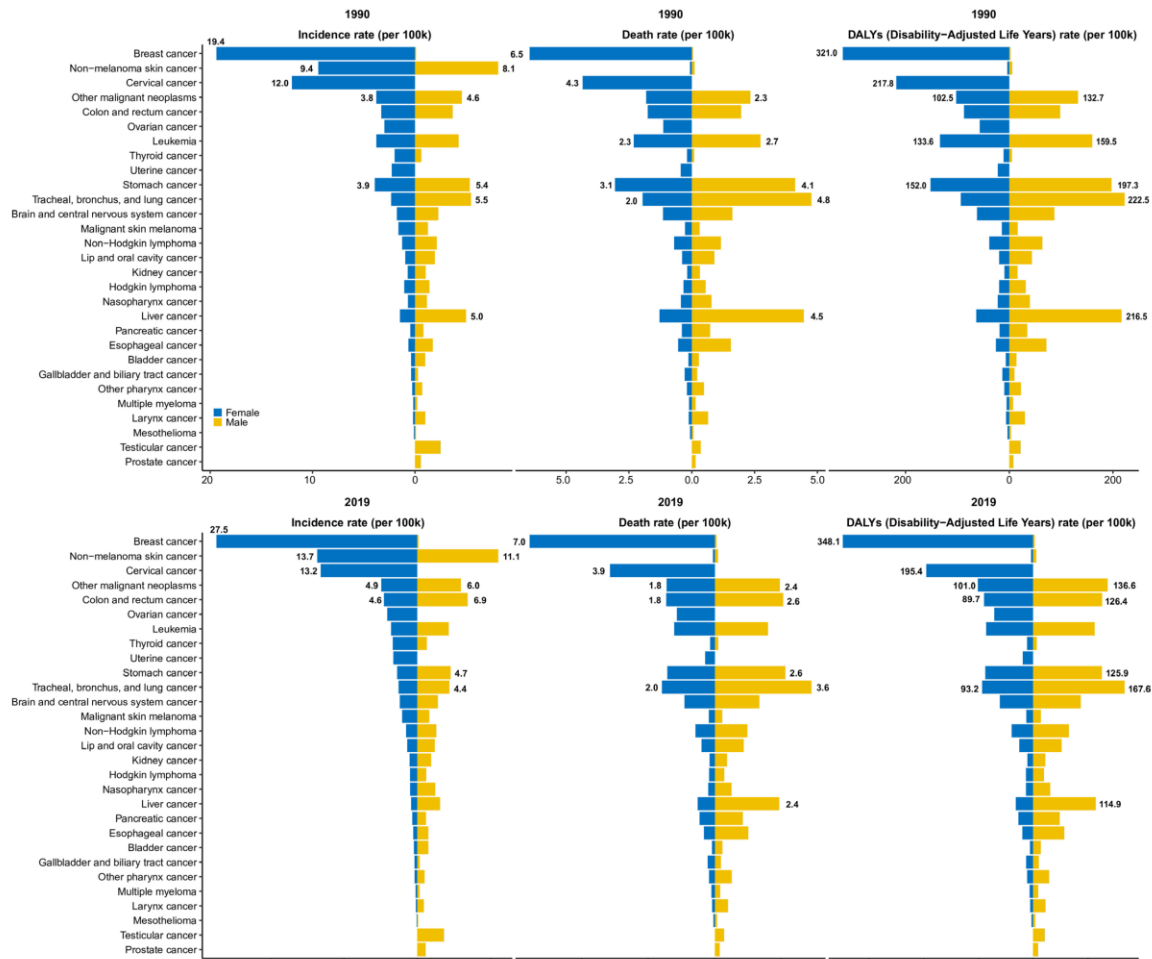
Trends in cancer incidence by age and sex, United States, 1998–2021

Increased incidence of 29 cancers in young people

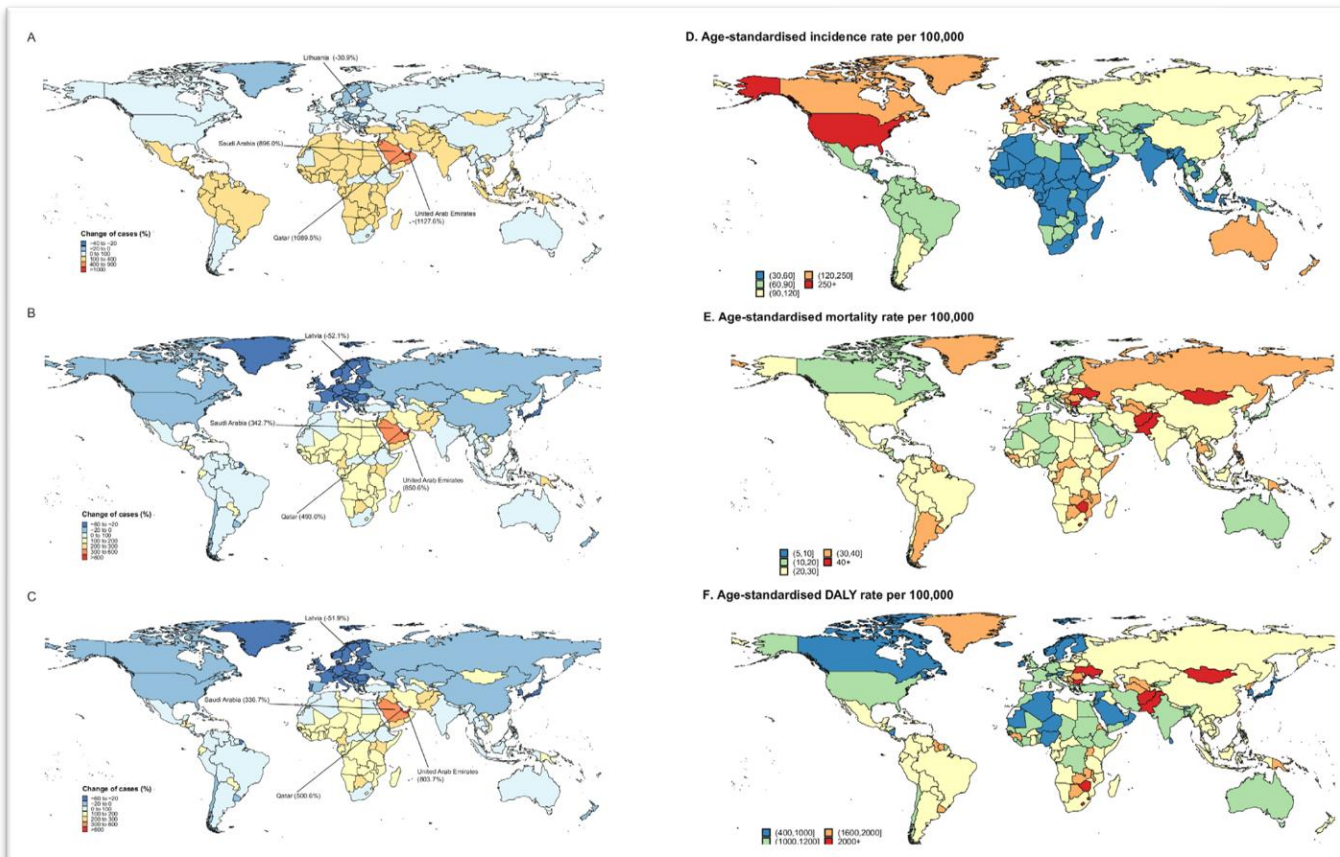


Between 1990 and 2019, the incidence of 29 cancers — including breast, lung and colorectal tumours — increased worldwide in people under the age of 50, particularly in women

Global trends in incidence, death, burden and risk factors of early-onset cancer from 1990 to 2019



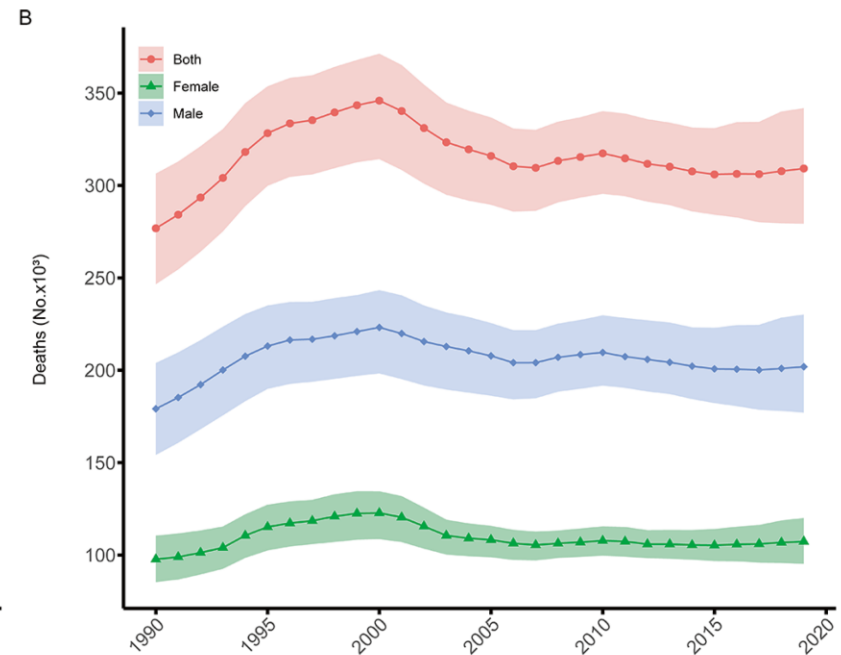
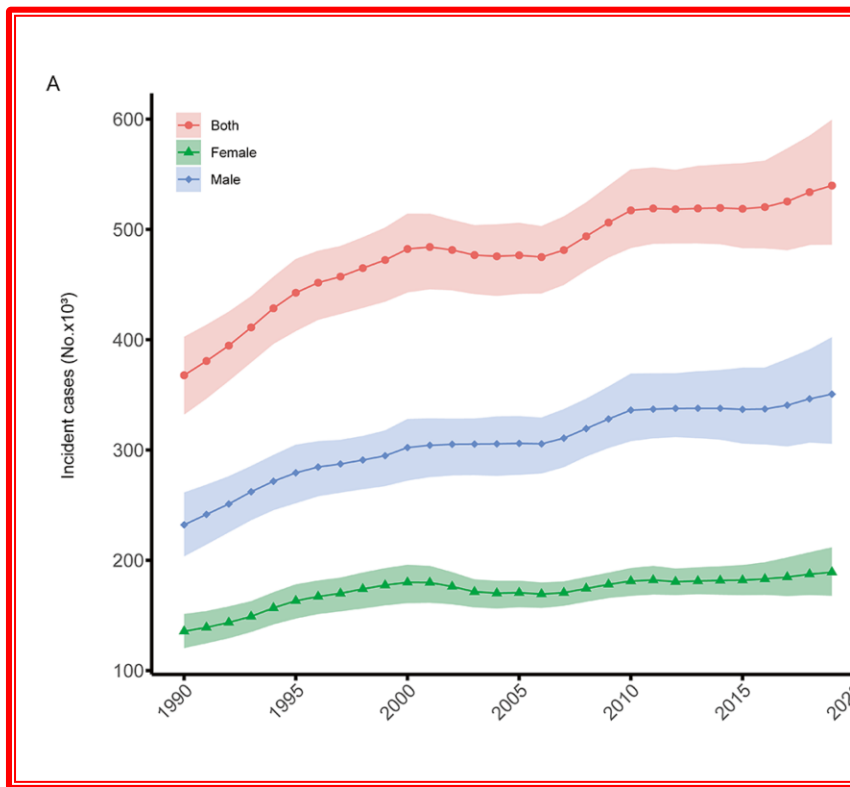
The global incidence, death and DALYs rates of 29 specified early-onset cancers in 1990 and 2019 by sex.



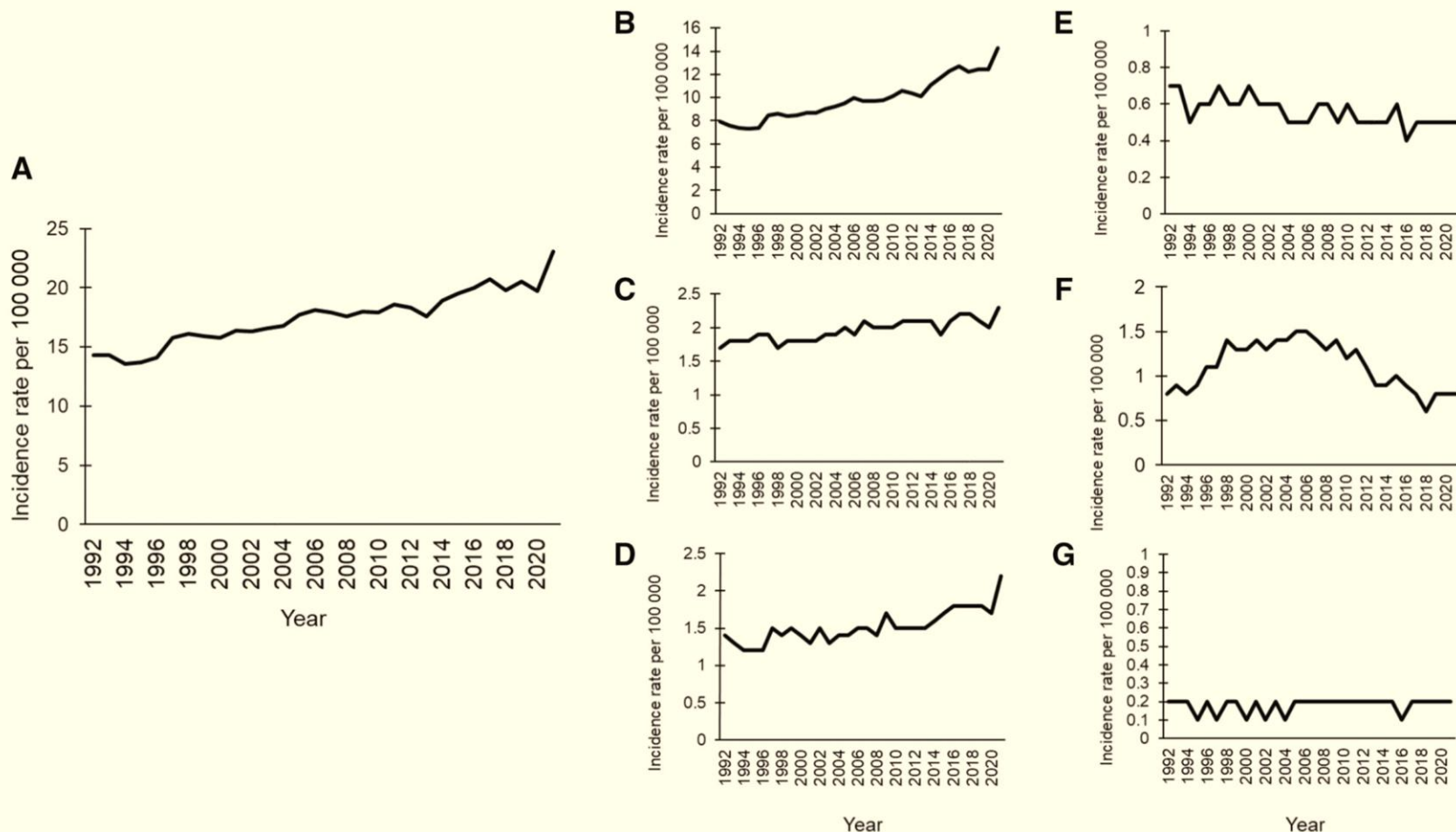
DALYs, disability-adjusted life years.

BMJ Oncology. 2023;2:e000049.
<https://doi.org/10.1136/bmjonc-2023-000049>

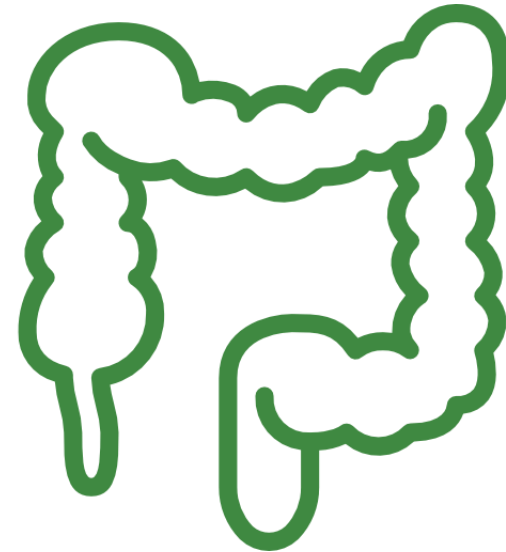
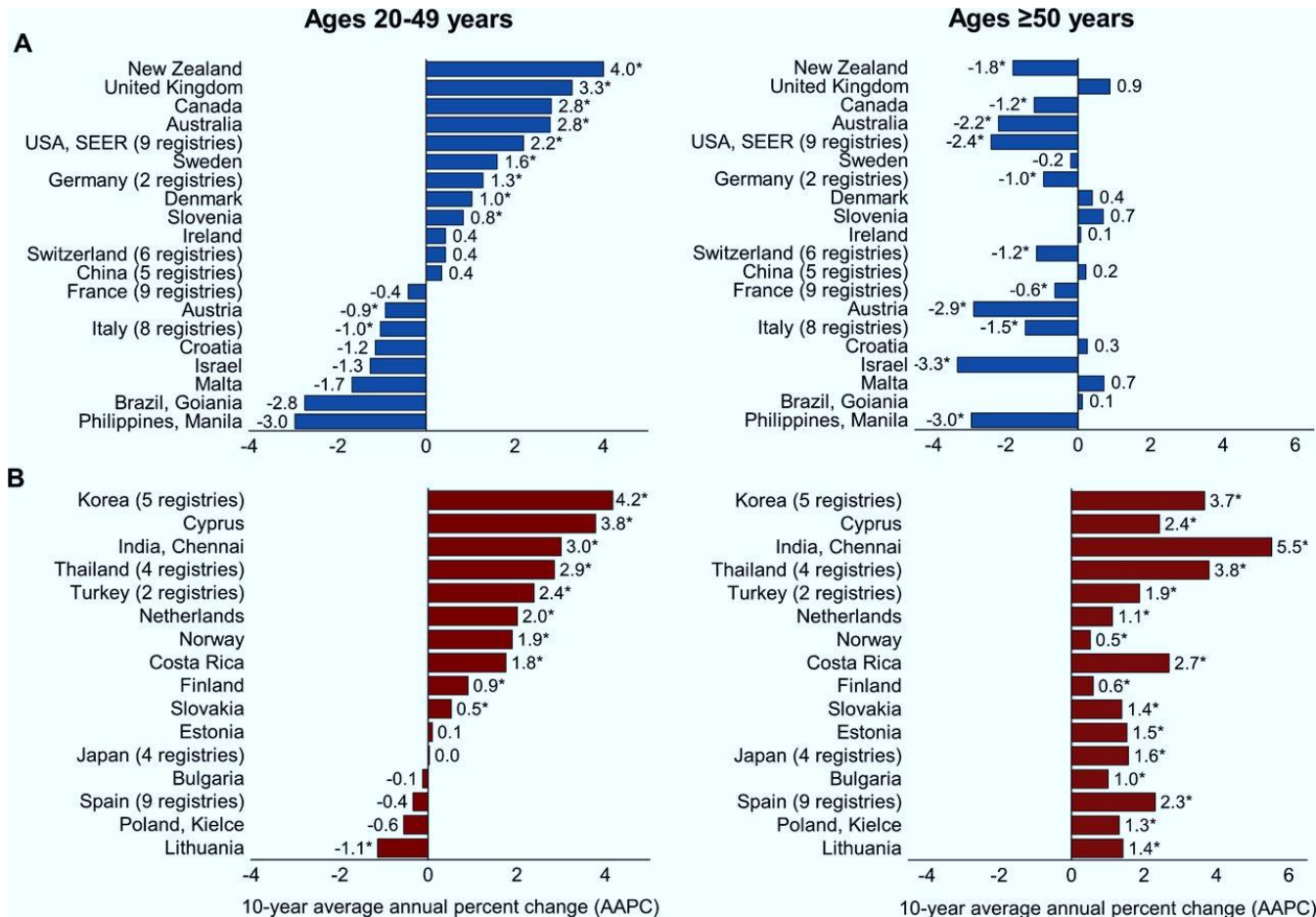
Total EO-GI cancer incidence and mortality absolute counts from 1990 to 2019



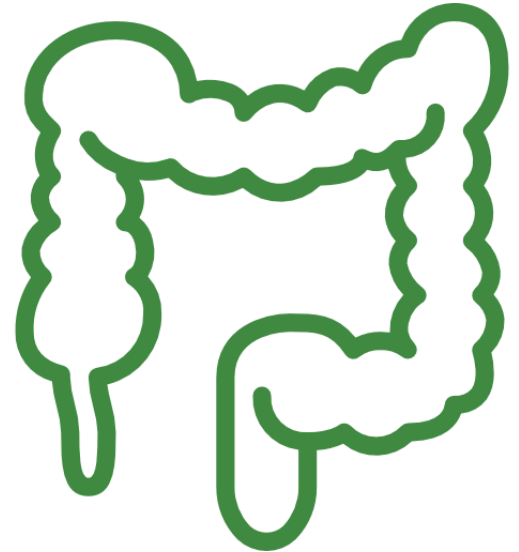
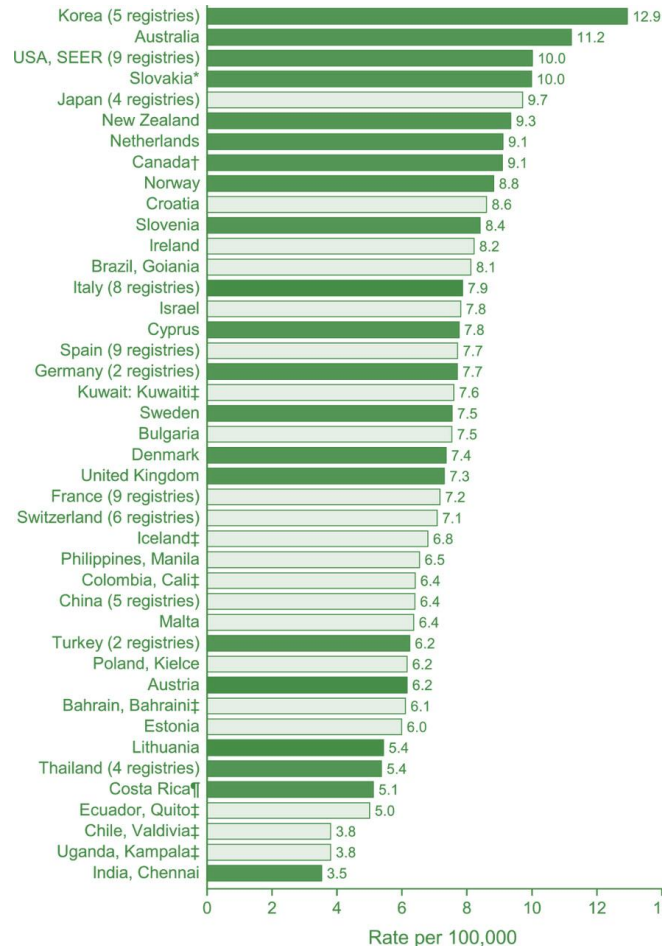
Increase in incidence rates of EOGLC, 1992-2021



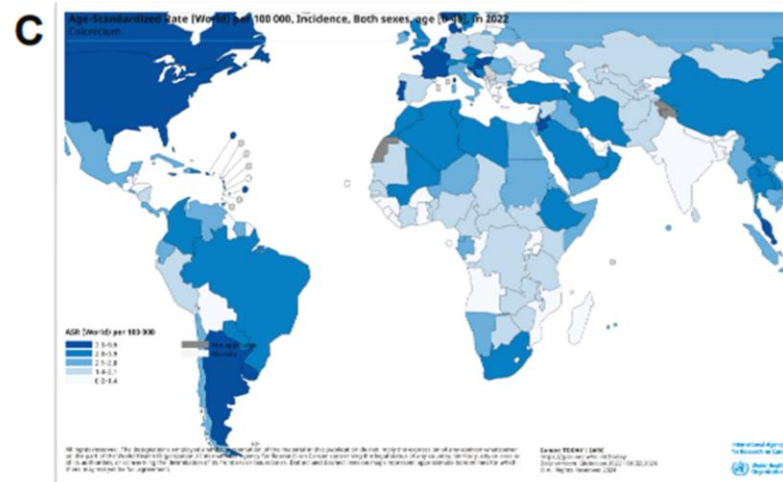
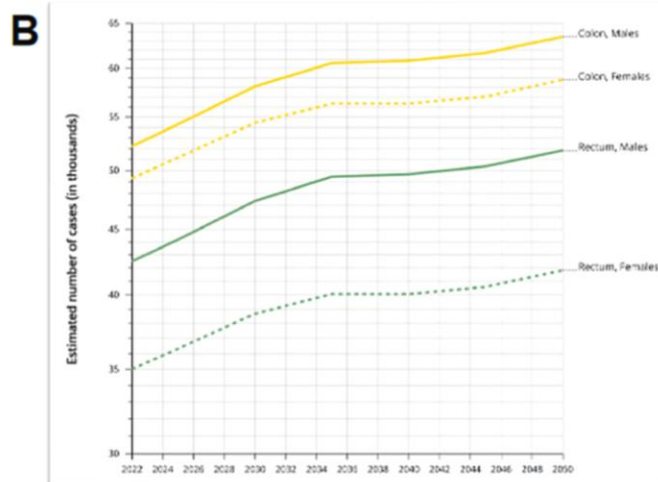
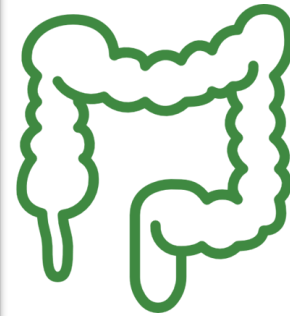
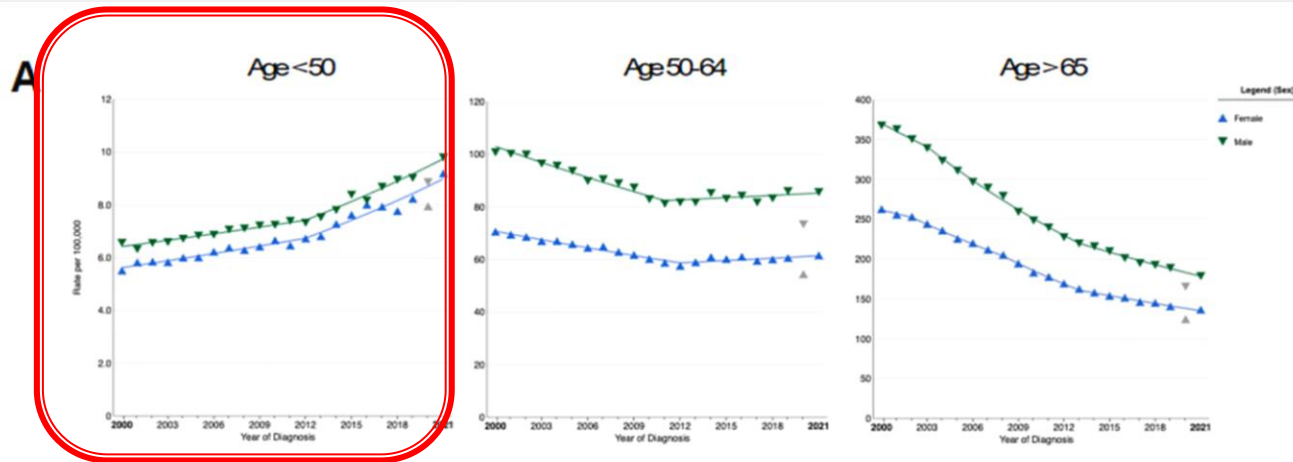
Average annual per cent change (AAPC) in colorectal cancer incidence by age during the most recent 10 years of available data.



Age-standardised incidence rate during 2008–2012 for colorectal cancer among adults ages 20–49 years.

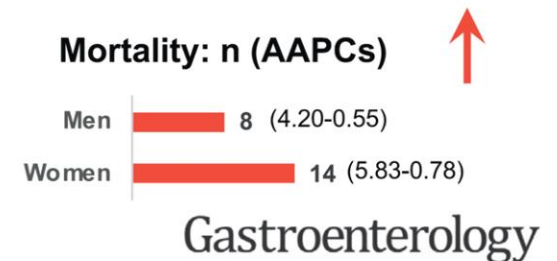
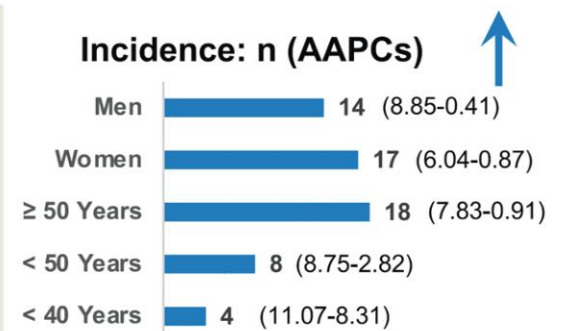
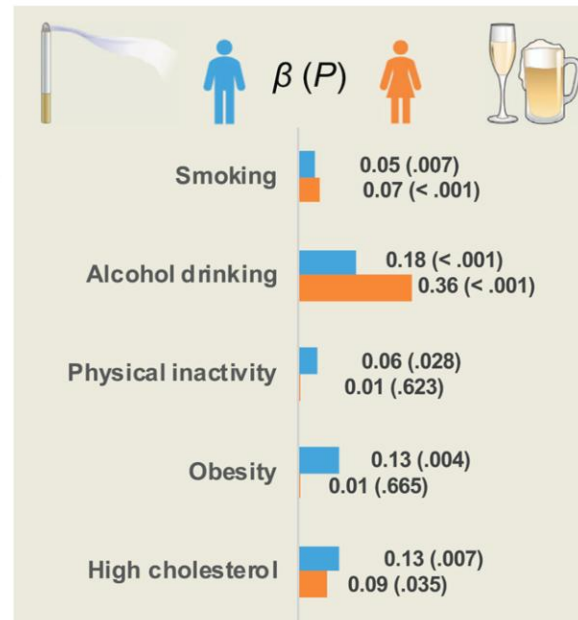
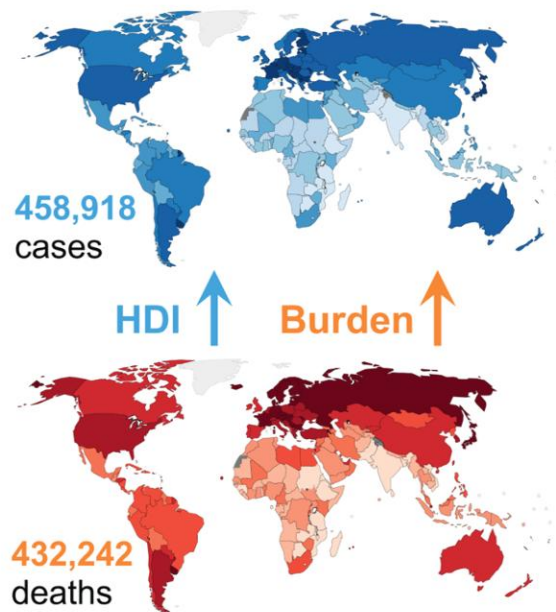


Epidemiology of EOCRC

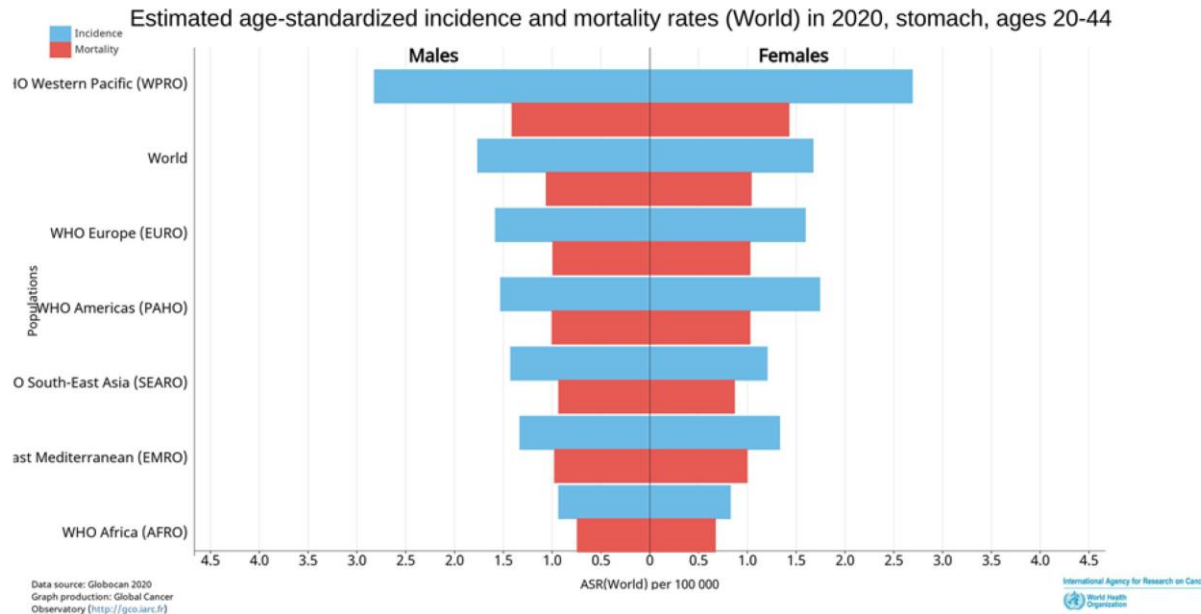


Worldwide Burden of Pancreatic Cancer

Worldwide Burden of, Risk Factors for, and Trends in Pancreatic Cancer



Global incidence and mortality trends of gastric cancer and predicted mortality of gastric cancer by 2035



In the past decade, the incidence and mortality of gastric cancer have shown a decreasing trend; however, there are still some countries showing an increasing trend, especially among populations younger than 45 years

Early-Onset Biliary Tract Cancer

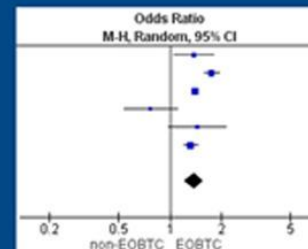
Early-Onset Biliary Tract Cancer (EO-BTC)

Epidemiology and Risk Factors

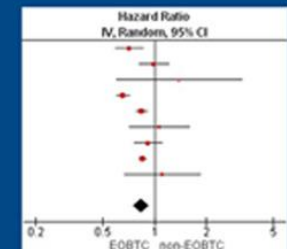


- EO-BTC: Definition remains heterogeneous across studies
- Epidemiologic trends suggest an increasing incidence of EO-iCCA
- Risk factors vary by anatomical subtype of EO-BTC

Treatment and Prognosis

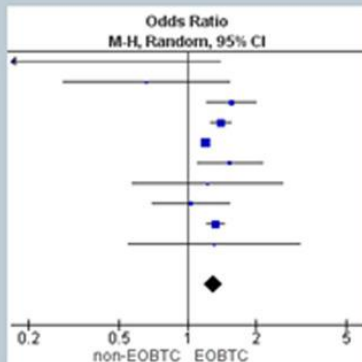


More surgery ↑

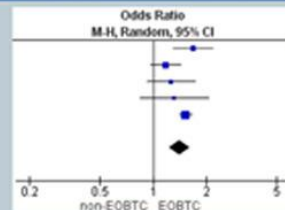


Better overall survival

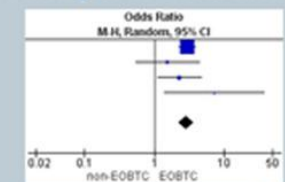
Clinicopathological and Molecular Features



Advanced stage at diagnosis ↑



Intrahepatic location ↑



FGFR2 fusions ↑

EO-BTC is a clinically and molecularly distinct subset within BTCs.

Prospective and age-stratified studies are needed to guide age-adapted detection and therapeutic strategies.



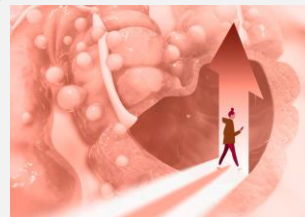
Early-Onset Gastrointestinal Cancers

Varies from country to country and cancer to cancer

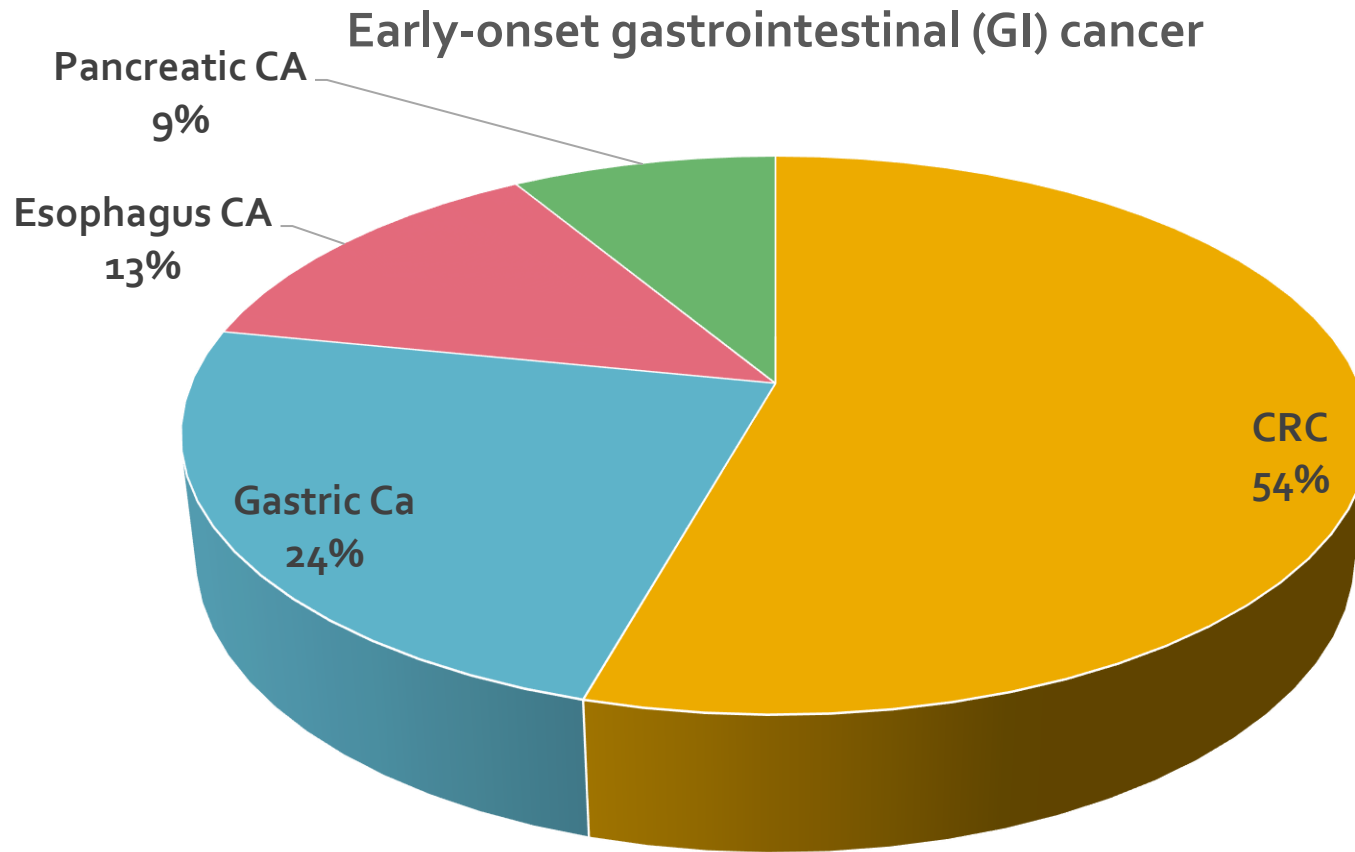
The incidence is increasing at an alarming rate globally

Women and in specific racial and ethnic groups

Birth cohort effect

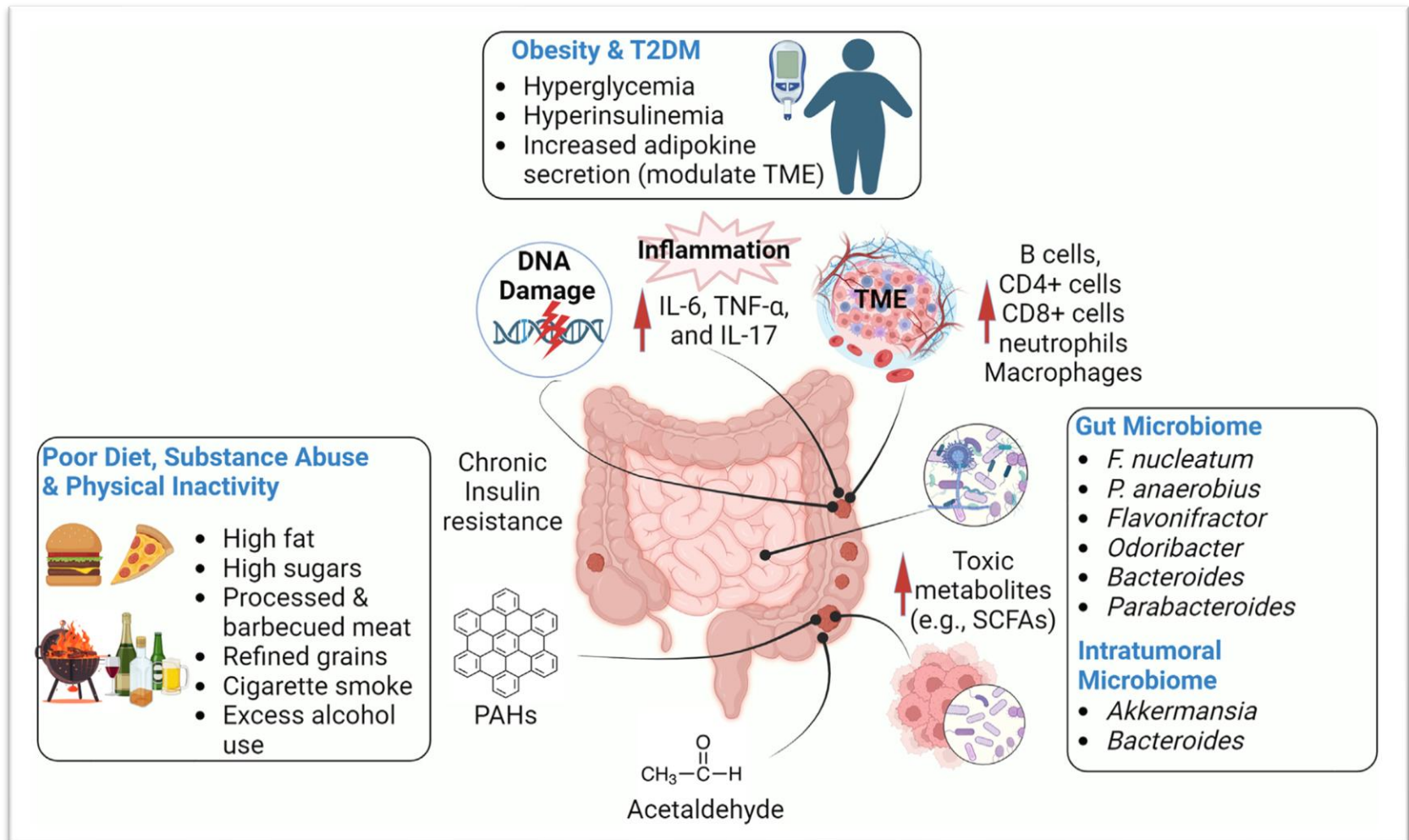


Worldwide early-onset GI cancers reported in 2022

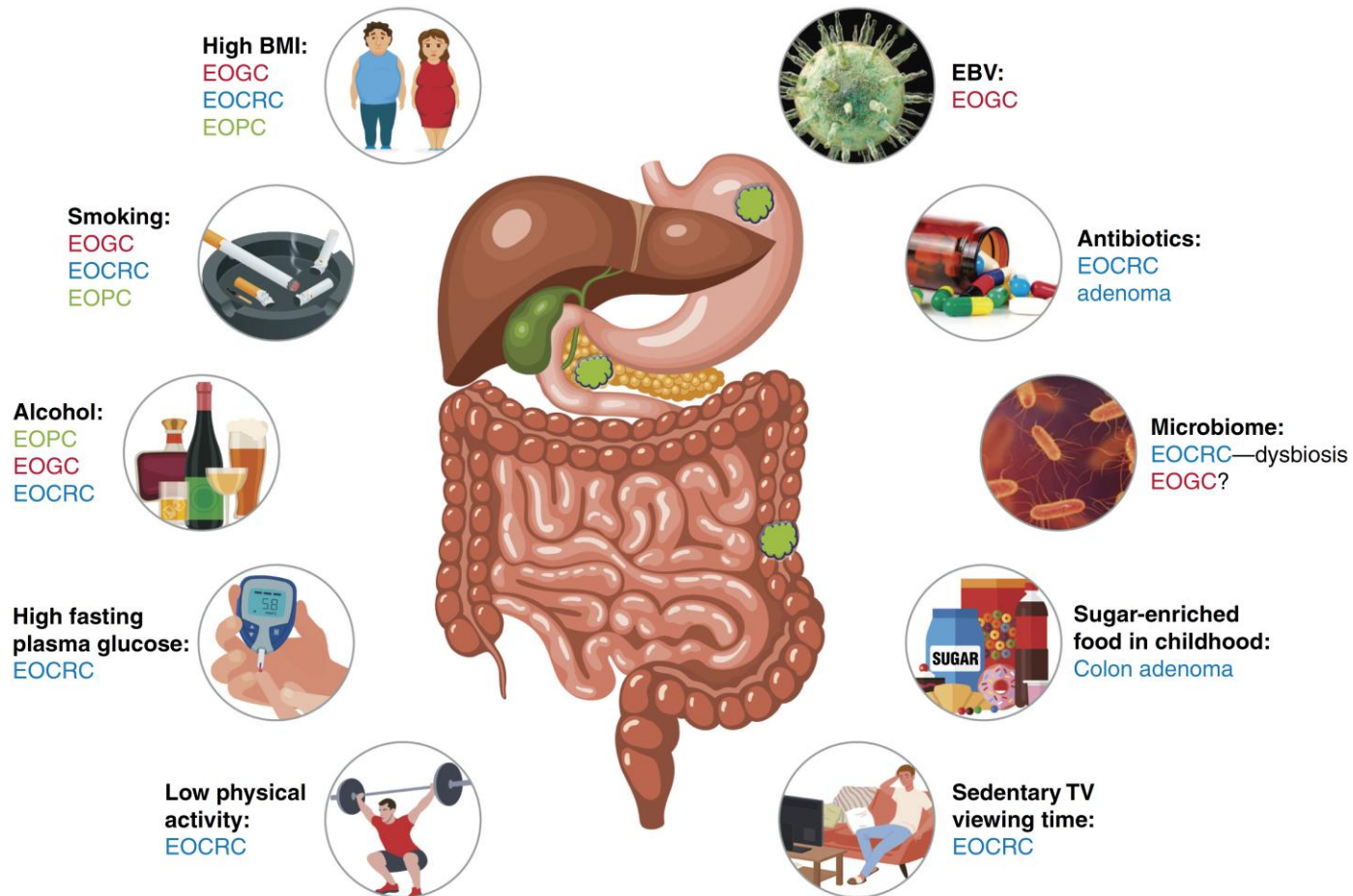


The incidence of early-onset GI cancer is rising globally, and early-onset GI cancers represent the most rapidly increasing early-onset cancer in the US.

Aetiology and Risk Factors



Risk Factors for EOGIC



Modifiable risk factors

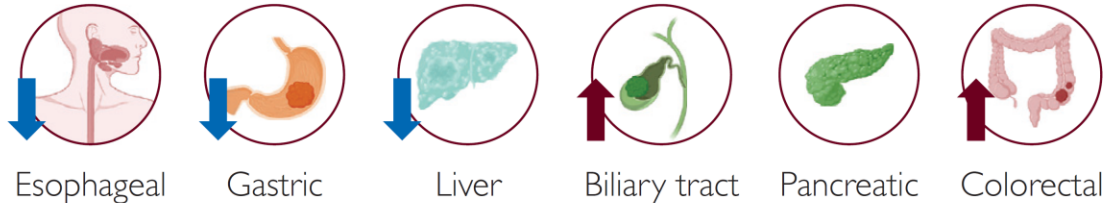
- Most early-onset GI cancers are associated with **modifiable risk factors** including:
 - Obesity,
 - Poor-quality diet (eg, sugar-sweetened beverages, ultraprocessed foods)
 - Sedentary lifestyle
 - Cigarette smoking
 - Alcohol consumption



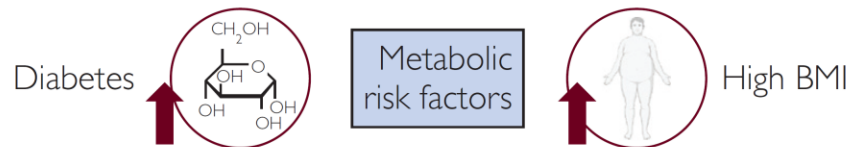
Metabolic Risk Factors

Increased incidence of **early-onset gastrointestinal cancer** and mortality from **metabolic risk factors**

Quantifying 6 types of early-onset (15-49 years) GI cancer using the global burden of disease study 2021



- 499,800 new cases, 285,900 deaths, and 14.01 million disability adjusted life years from early-onset GI cancer in 2021
- Early-onset GI cancer accounted for 9.51% of the overall GI cancer incidence
- From 2000 to 2021, early-onset colorectal and biliary tract cancer increased in incidence rates



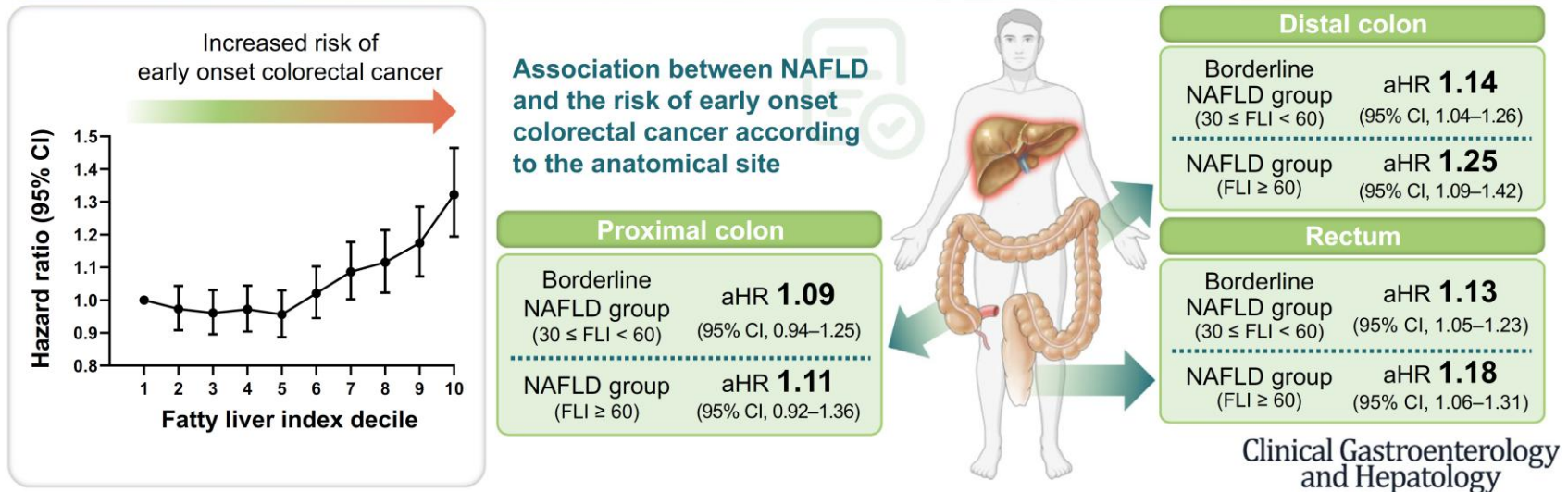
Death and disability-adjusted life year rate of early-onset cancer attributable to metabolic risk factors increased in all types

Danpanichkul et al

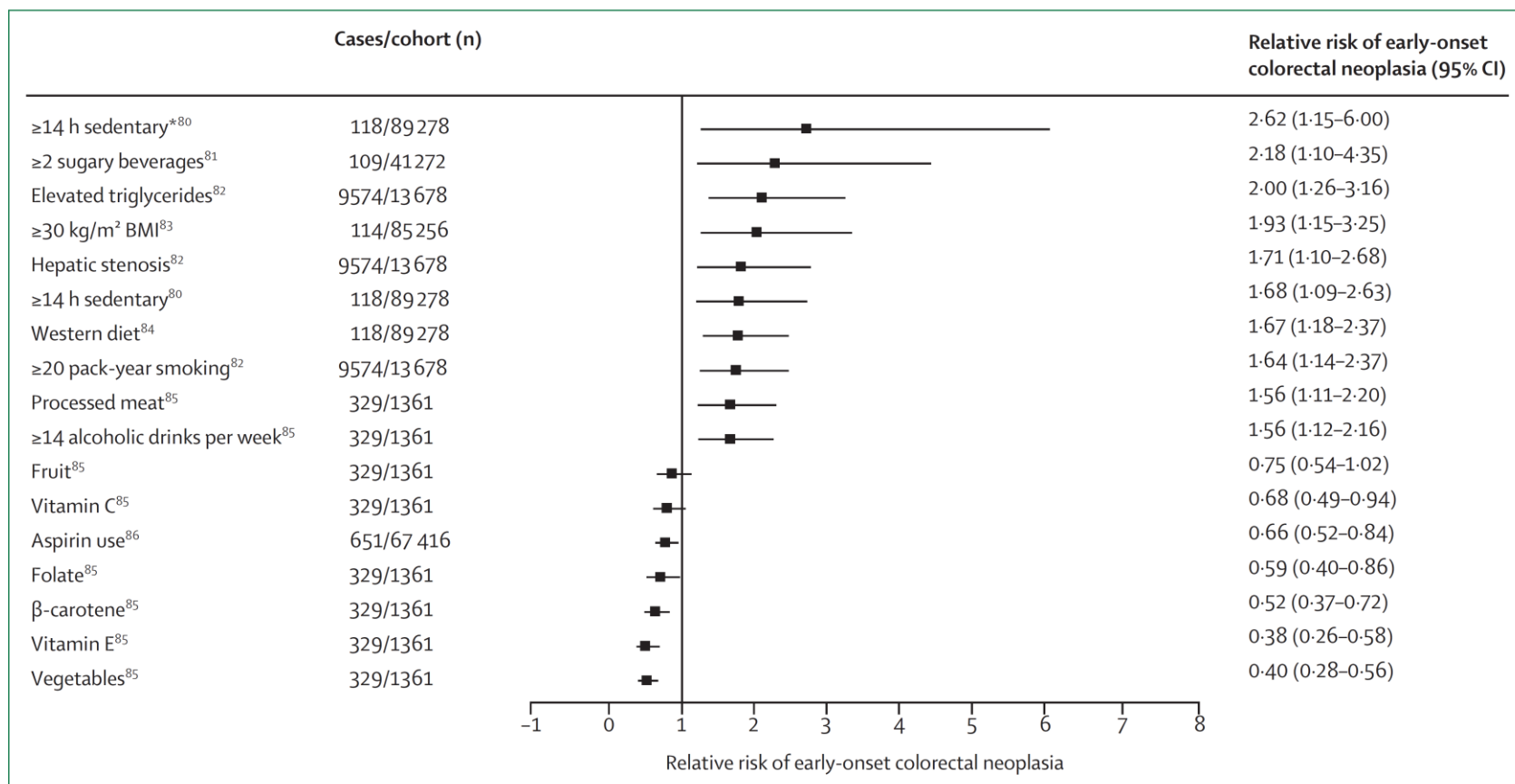
Association Between Nonalcoholic Fatty Liver Disease and Risk of Early-onset Colorectal Cancer

NAFLD, including borderline cases, is significantly associated with an increased risk of early-onset CRC, particularly in the distal colon and rectum

Nonalcoholic fatty liver disease and early onset colorectal cancer - A Nationwide-population based cohort study in Korea -

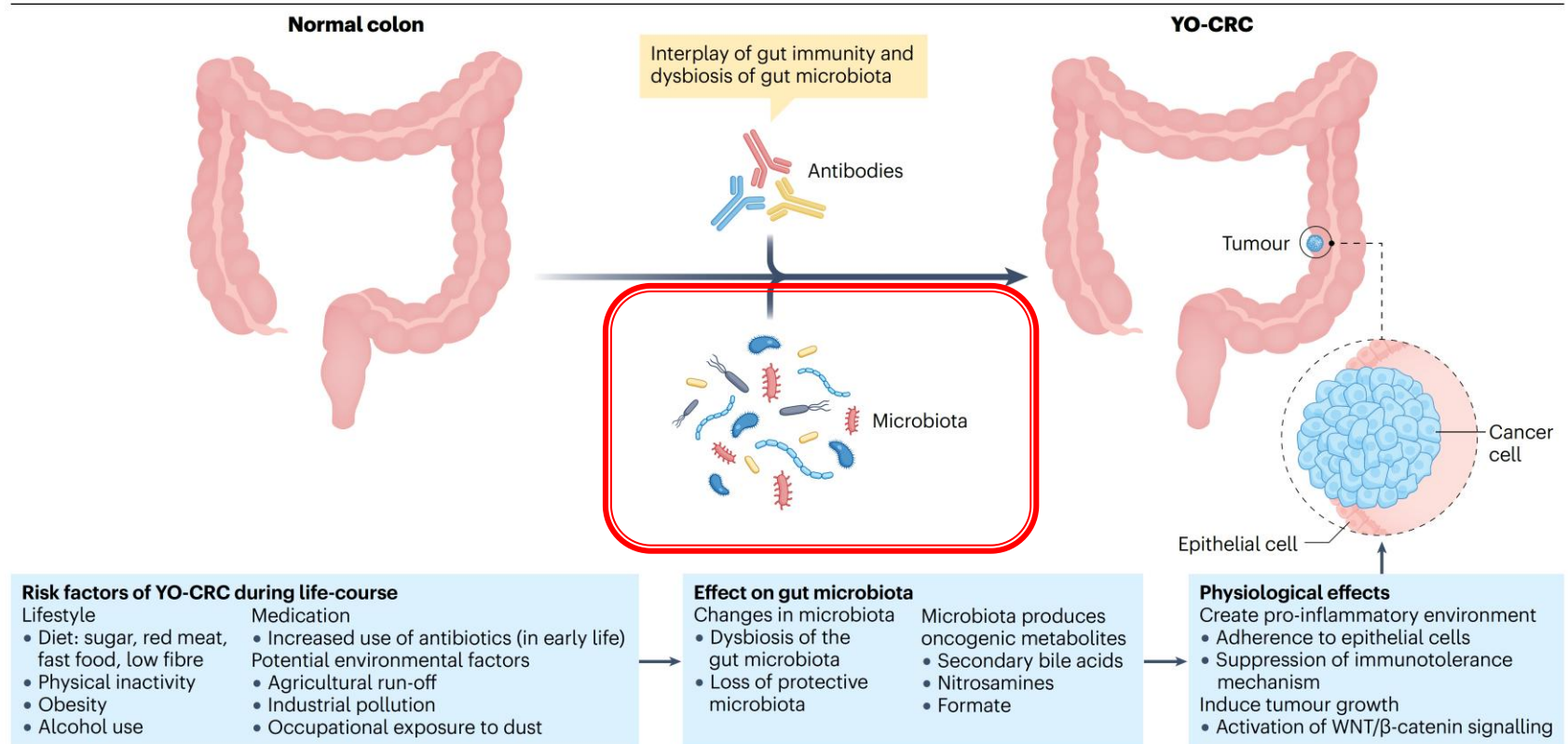


Diet and lifestyle factors associated with early-onset colorectal cancer or neoplasia



BMI=body-mass index. Western diet=diet high in ultra-processed foods. *Rectal cancer

Pathogenetic mechanism of sporadic EO-CRC



Spaander, M.C.W et al. Nat Rev Dis Primers 9, 21 (2023).
<https://doi.org/10.1038/s41572-023-00432-7>

Early-onset vs. Late-onset cancers

Early-onset colorectal cancer

age <50

Features that are more frequent in EOCRC

Family history and microsatellite instability

Metastatic, late-stage diagnoses

Poor differentiation, LINE-1 (long interspersed nuclear element 1) hypomethylation

Clinical presentation with a predilection for the left side of the colon and rectum

Features that are frequent in both EOCRC and LOCRC

Metabolic syndrome

Consumption of sugar-sweetened beverages, red and processed meat

Western pattern diet

Obesity and prolonged sedentary behavior

Later-onset colorectal cancer

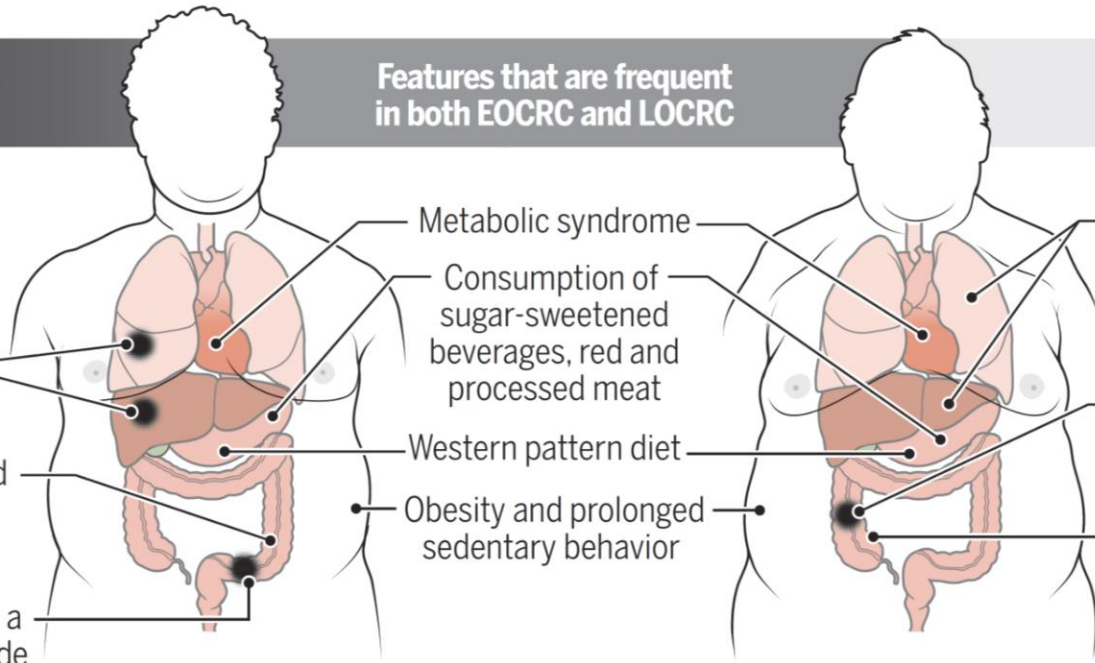
age ≥50

Features that are more frequent in LOCRC

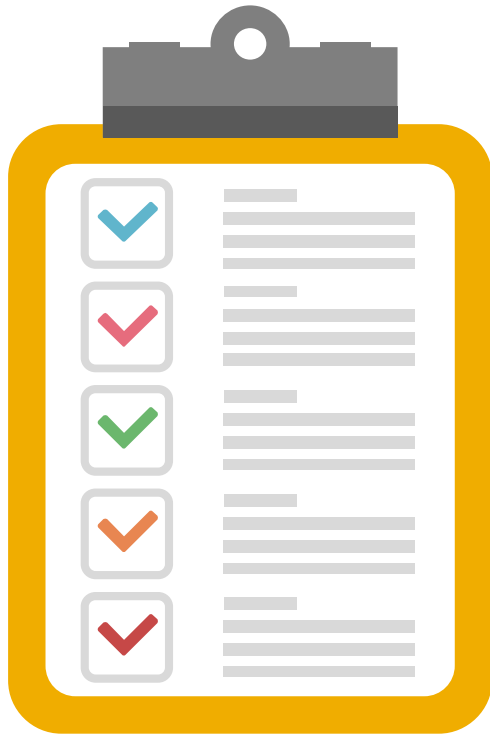
Earlier detection (screen-detected) and lower likelihood of metastases

Tumor adenomatous polyposis coli (APC) and BRAF mutations

Clinical presentation with a predilection for the right side of the colon



Unique Clinical and Molecular Characteristics



- 1 Aggressive Presentation
- 2 Distinct Molecular Profiles
- 3 Aggressive Treatment
- 4 Inconsistent Survival
- 5 Disparities

Implications of Early-onset GI Cancers

Cancer Screening
Policies

Patient Care and
Survivorship



Clinical Practice and Diagnosis

Medical Research

1. Implication for Clinical Practice and Diagnosis

Increased clinical suspicion of GI cancer in younger patients.

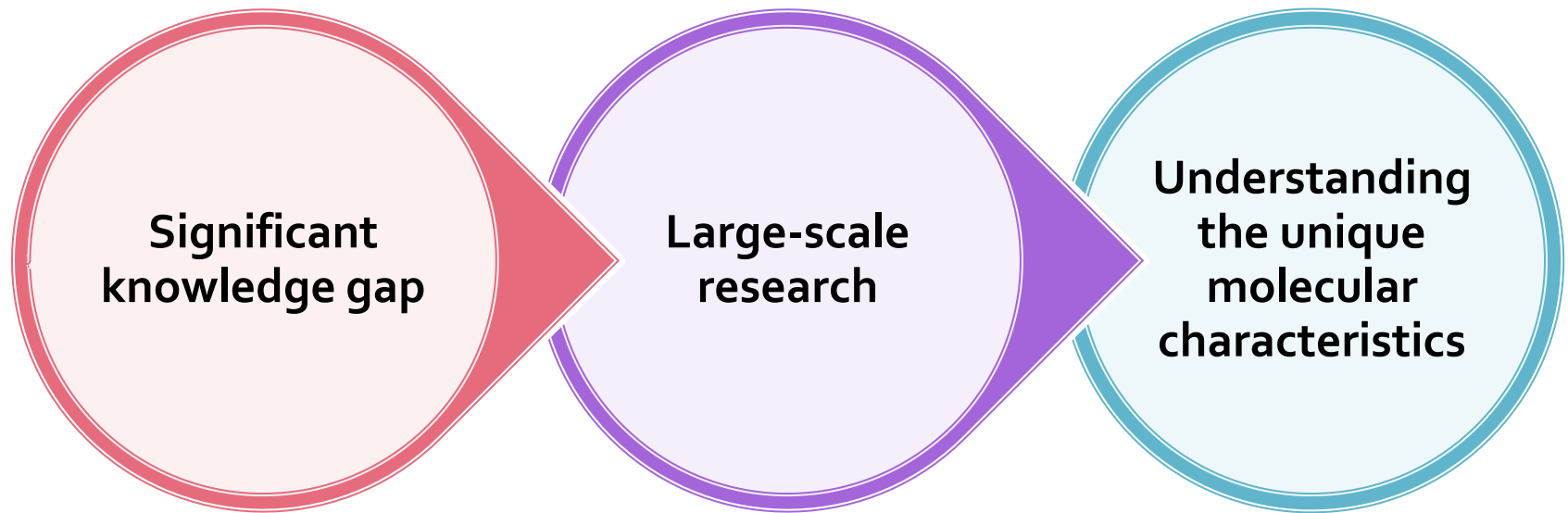
Healthcare providers must be more vigilant and consider GI cancer in their differential diagnosis for younger symptomatic patients

2. Implication for Cancer Screening Policies

Further adjustments to screening strategies

Developing effective, non-invasive screening tools.

3. Implication for Medical Research



4. Implication for Patient Care and Survivorship

Holistic

Multidisciplinary

Extending beyond medical treatment to include:

- **Fertility Preservation**
- **Psychosocial and Financial Support**
- **Genetic Counseling**

Future Directions

01

**Improved
Screening**

02

**Dedicated
Research**

03

**Specialized
Care**

Roadmap of strategies to reduce the public health burden of early-onset CRC

Theme	Strategy
Education	Educate physicians, other health-care workers and the general public in order to raise awareness of early-onset colorectal cancer
Screening	<ul style="list-style-type: none">▪ Expand the use of genetic testing among individuals with a family history▪ Conduct research to create and refine prediction models▪ Evaluate the available evidence and screening guidelines▪ Develop personalized screening strategies
Aetiological research	Utilize existing resources to study the aetiologies of early-onset CRC; design additional studies
Clinical care and research	Set up specialized centres, units and/or clinics focused on early-onset CRC in order to deliver optimal care; conduct clinical trials

