

CONGRESSO REGIONALE CALABRIA SID - AMD 2022



T-HOTEL LAMEZIA TERME
25-26 NOVEMBRE 2022

Diabete e cancro

Domenico Voce
Diabetologia Crotone



CONGRESSO REGIONALE CALABRIA SID - AMD 2022



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25-26 NOVEMBRE 2022





diabetes and covid-19



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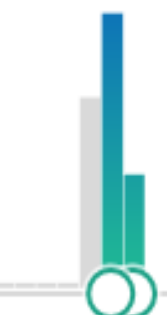
4,774 results

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RESULTS BY YEAR



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diabetes and cancer



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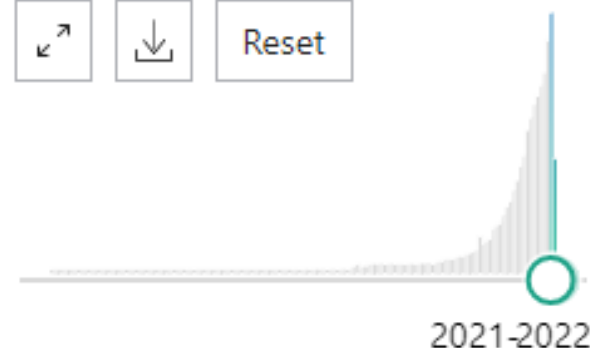
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7,695 results

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RESULTS BY YEAR



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☐ Obesity, **Diabetes**, and Increased **Cancer** Progression.

1 Kim DS, Scherer PE.

Cite Diabetes Metab J. 2021 Nov;45(6):799-812. doi: 10.4093/dmj.2021.0077. Epub 2021 Nov 22.
PMID: 34847640 [Free PMC article.](#) [Review.](#)

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Many observations suggest that obesity and **diabetes** are associated with an increased risk of developing several types of **cancers**, including liver, pancreatic, endometrial, colorectal, and post-menopausal breast **cancer**. ...Here, we highlight the cellular and m ...

TEXT AVAILABILITY



diabetes and stroke



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3,228 results

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RESULTS BY YEAR



2021-2022

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Diabetes, stroke, and neuroresilience: looking beyond hyperglycemia.

1

Krinock MJ, Singhal NS.

Cite

Ann N Y Acad Sci. 2021 Jul;1495(1):78-98. doi: 10.1111/nyas.14583. Epub 2021 Feb 26.

PMID: 33638222 Review.

Share

Additional research into clinical therapies directed at **diabetic** pathophysiological processes to prevent **stroke** and improve outcome for **diabetic stroke** survivors is necessary. ...Our review summarizes the underpinnings of **stroke** risk in **diabe** ...

TEXT AVAILABILITY

Number of people with diabetes worldwide and per IDF Region in 2021–2045 (20–79 years)

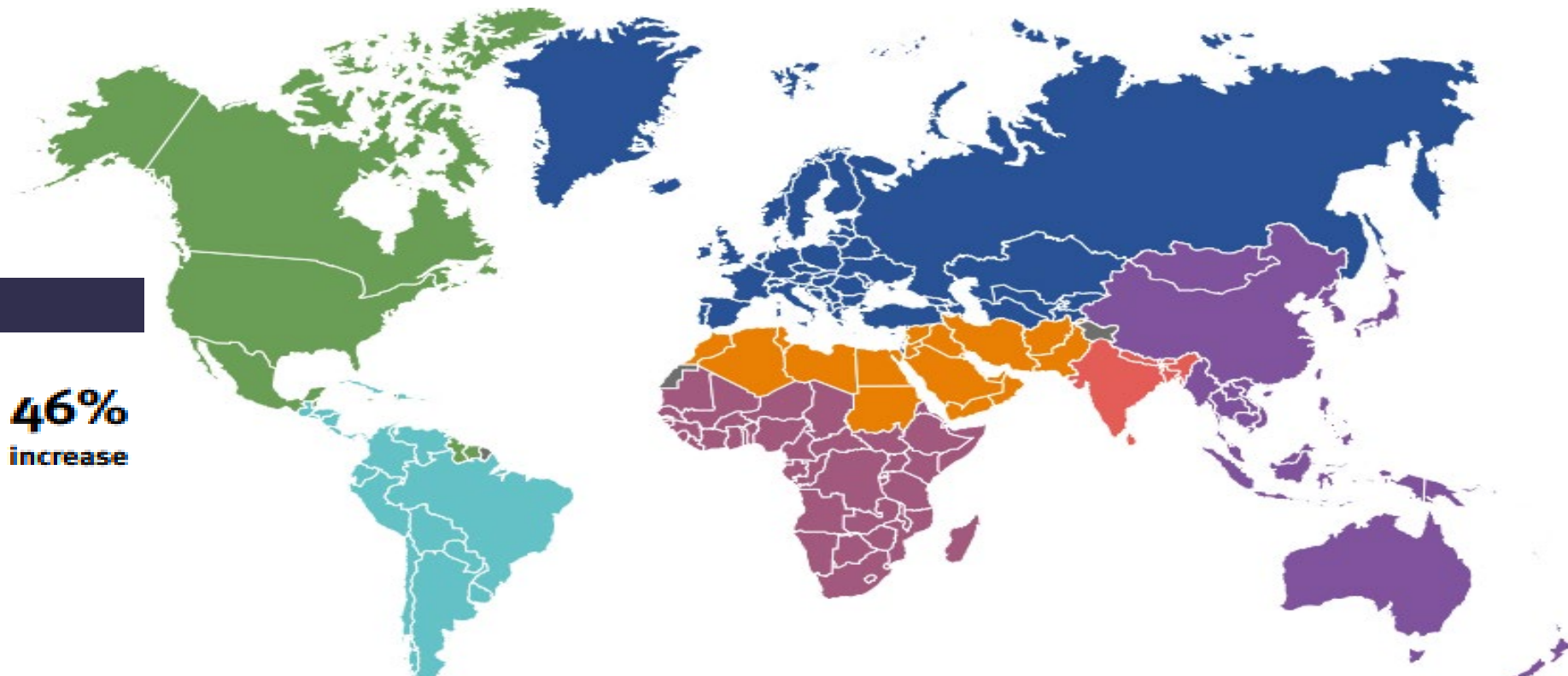
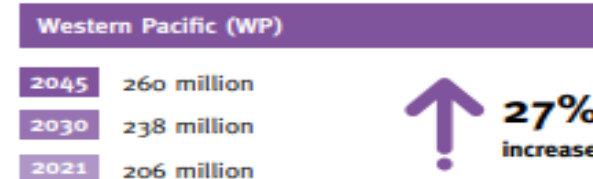
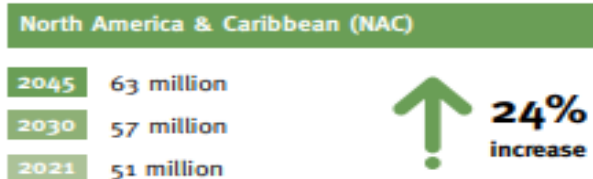
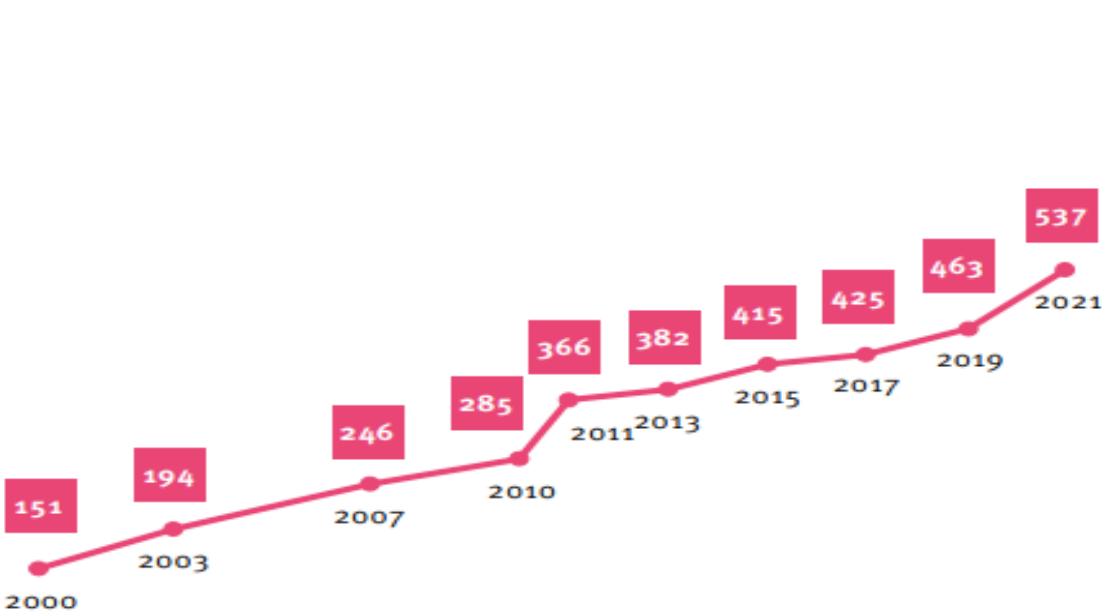


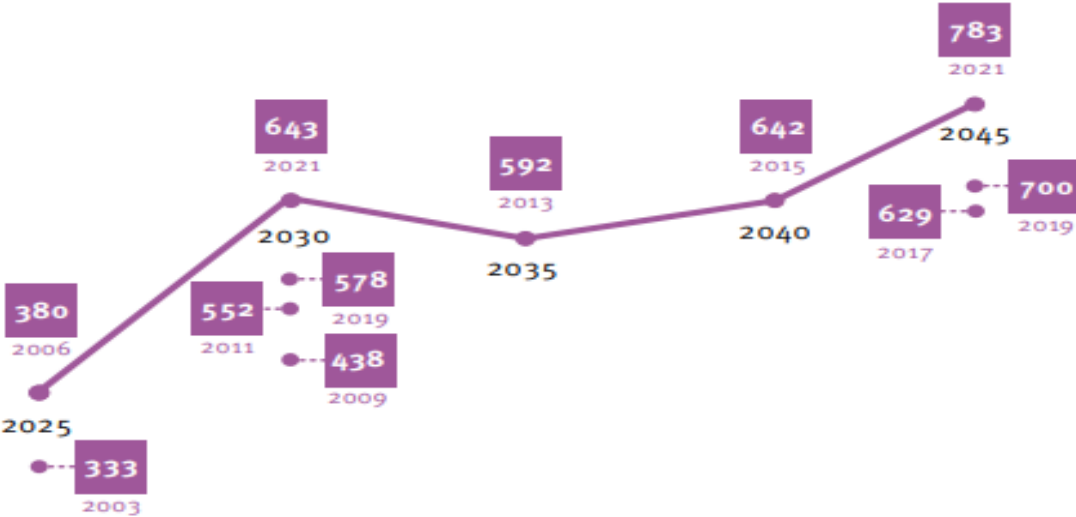
Figure 1 Estimates and projections of the global prevalence of diabetes in the 20–79 year age group in millions (IDF Diabetes Atlas editions 1st to 10th)

Estimates of the global prevalence of diabetes in the 20–79 year age group (millions)



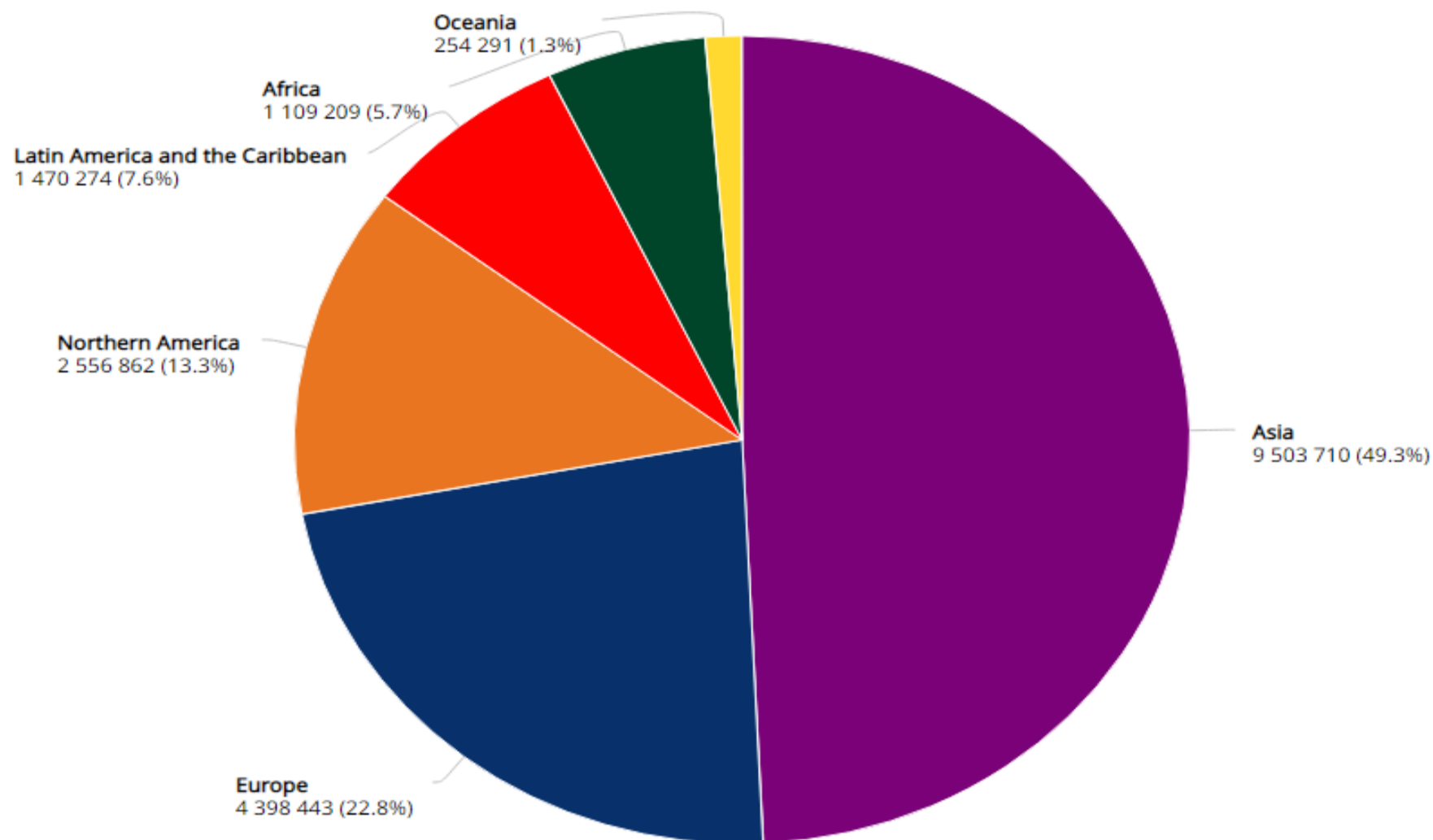
Key
151
Number of people with diabetes in millions

Projections of the global prevalence of diabetes in the 20–79 year age group (millions)



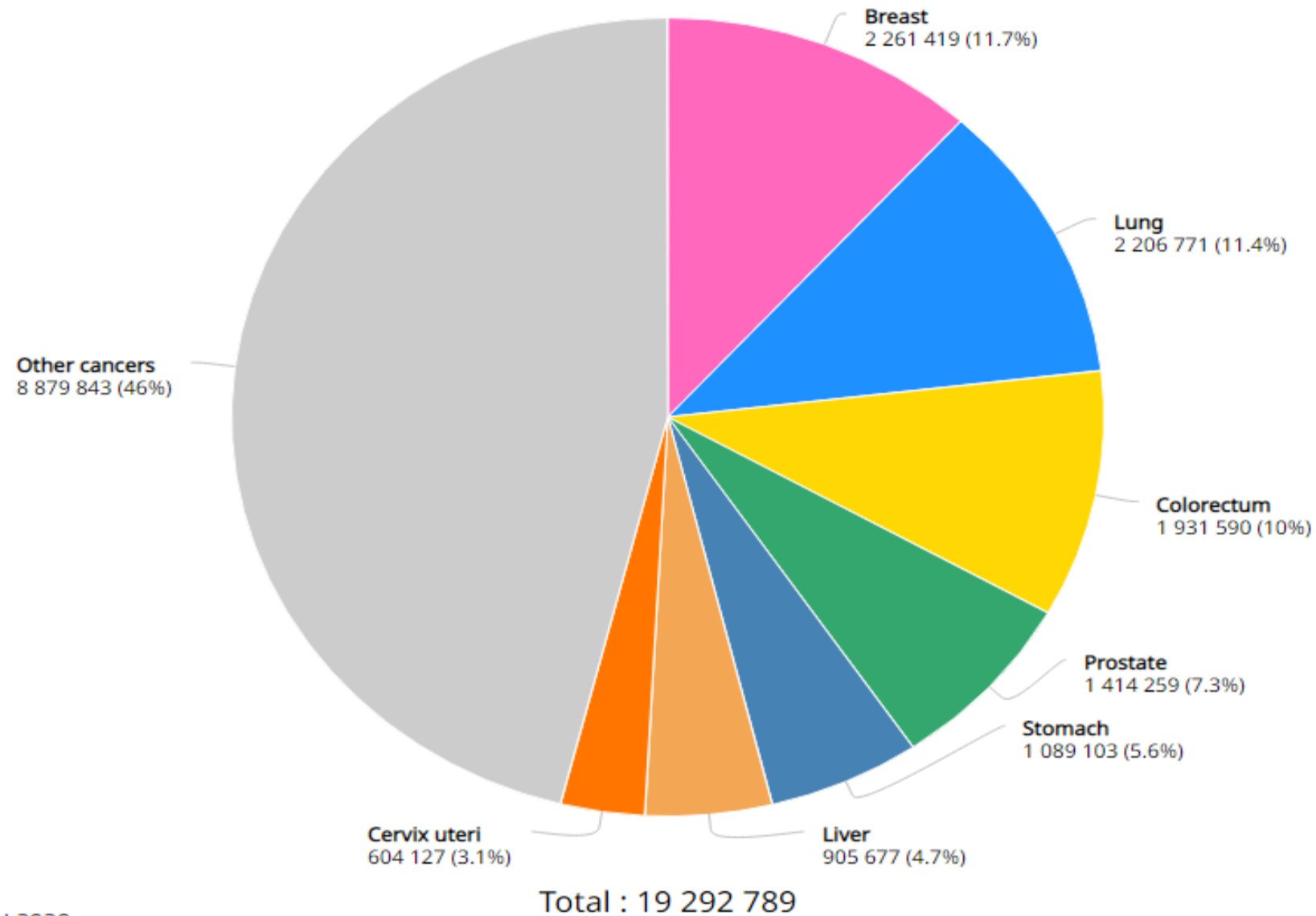
Key
333
2003
Projection in millions
Year projection made

Estimated number of new cases in 2020, all cancers, both sexes, all ages

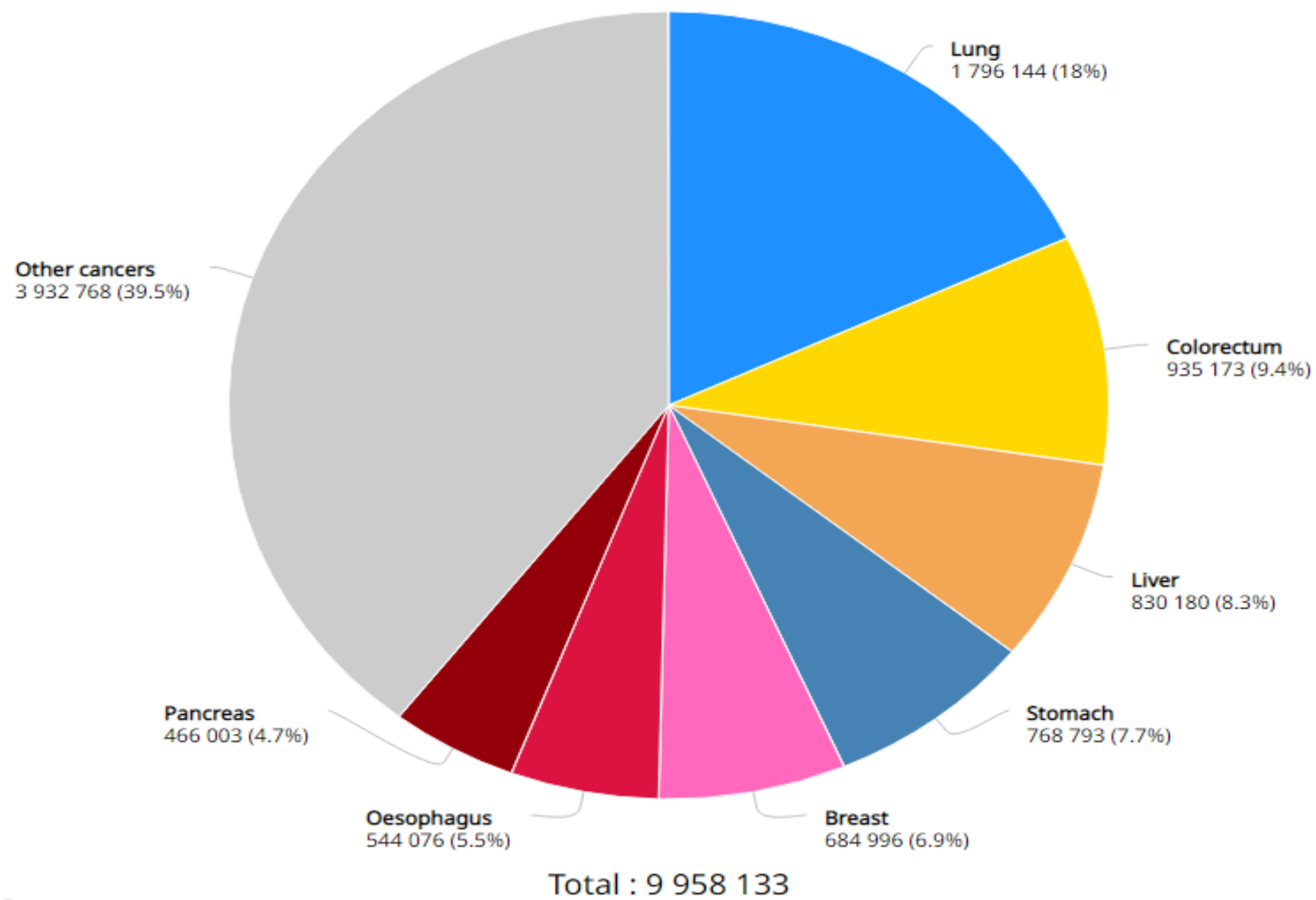


Total : 19 292 789

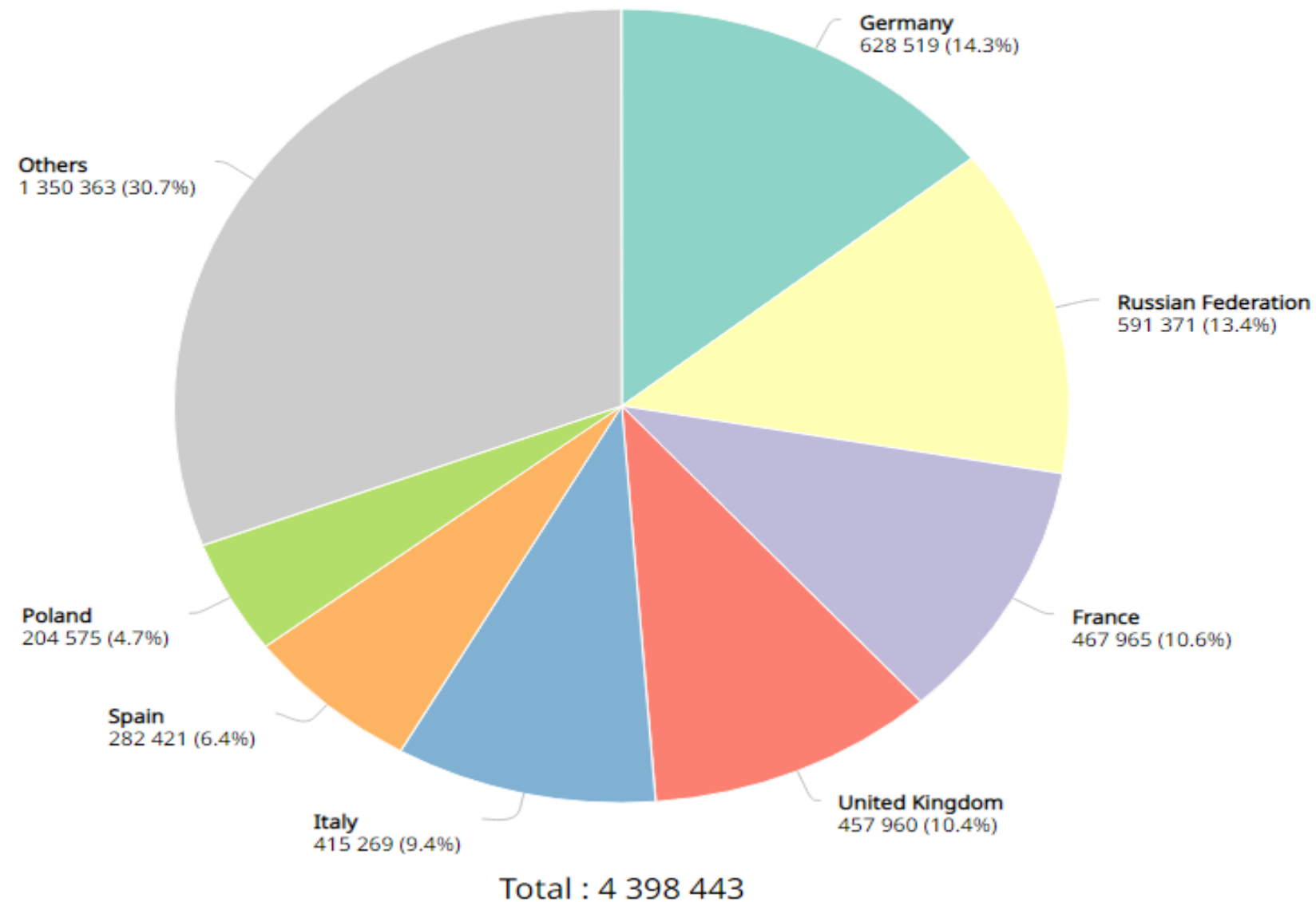
Estimated number of new cases in 2020, worldwide, both sexes, all ages



Estimated number of deaths in 2020, worldwide, both sexes, all ages



Estimated number of new cases in 2020, all cancers, both sexes, all ages



Estimated number of new cases from 2020 to 2040, Both sexes, age [0-85+]

All cancers

Africa + Latin America and Caribbean + Northern America + Europe + Oceania + Asia

2020



19.3M

2040



28.9M



= 1 000 000

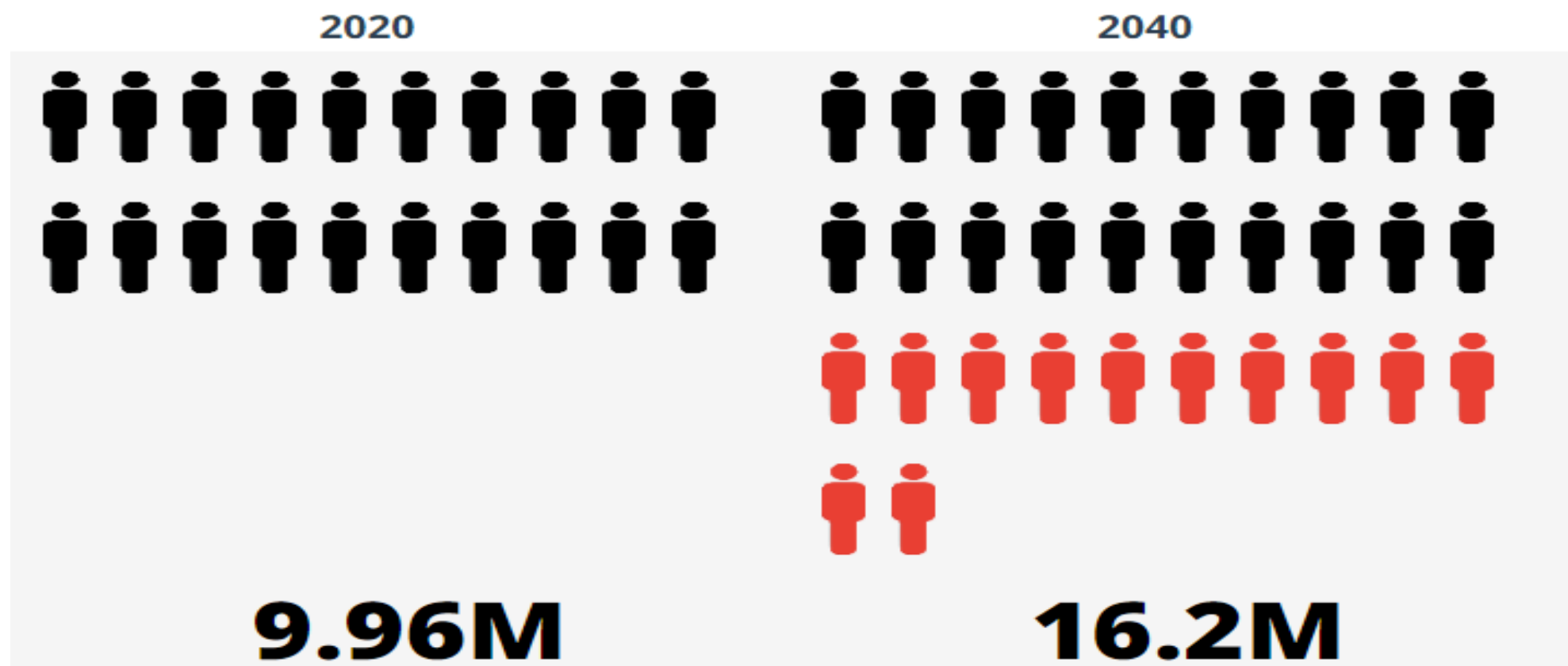


Demographic changes

Estimated number of deaths from 2020 to 2040, Both sexes, age [0-85+]

All cancers

Africa + Latin America and Caribbean + Northern America + Europe + Oceania + Asia



= 500 000



Demographic changes

Leading causes 2016

1 Ischaemic heart disease
2 Stroke
3 Lower respiratory infections
4 Diarrhoeal diseases
5 Road injuries
6 Malaria
7 Neonatal preterm birth
8 HIV/AIDS
9 COPD
10 Neonatal encephalopathy
11 Tuberculosis
12 Congenital defects
13 Lung cancer
14 Self-harm
15 Diabetes
16 Chronic kidney disease
17 Other neonatal
18 Alzheimer's disease
19 Neonatal sepsis
20 Liver cancer

25 Falls

26 Colorectal cancer

28 Hypertensive heart disease

29 Breast cancer

Leading causes 2040

1 Ischaemic heart disease
2 Stroke
3 Lower respiratory infections
4 COPD
5 Chronic kidney disease
6 Alzheimer's disease
7 Diabetes
8 Road injuries
9 Lung cancer
10 Diarrhoeal diseases
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17 Congenital defects
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19 Breast cancer
20 Falls

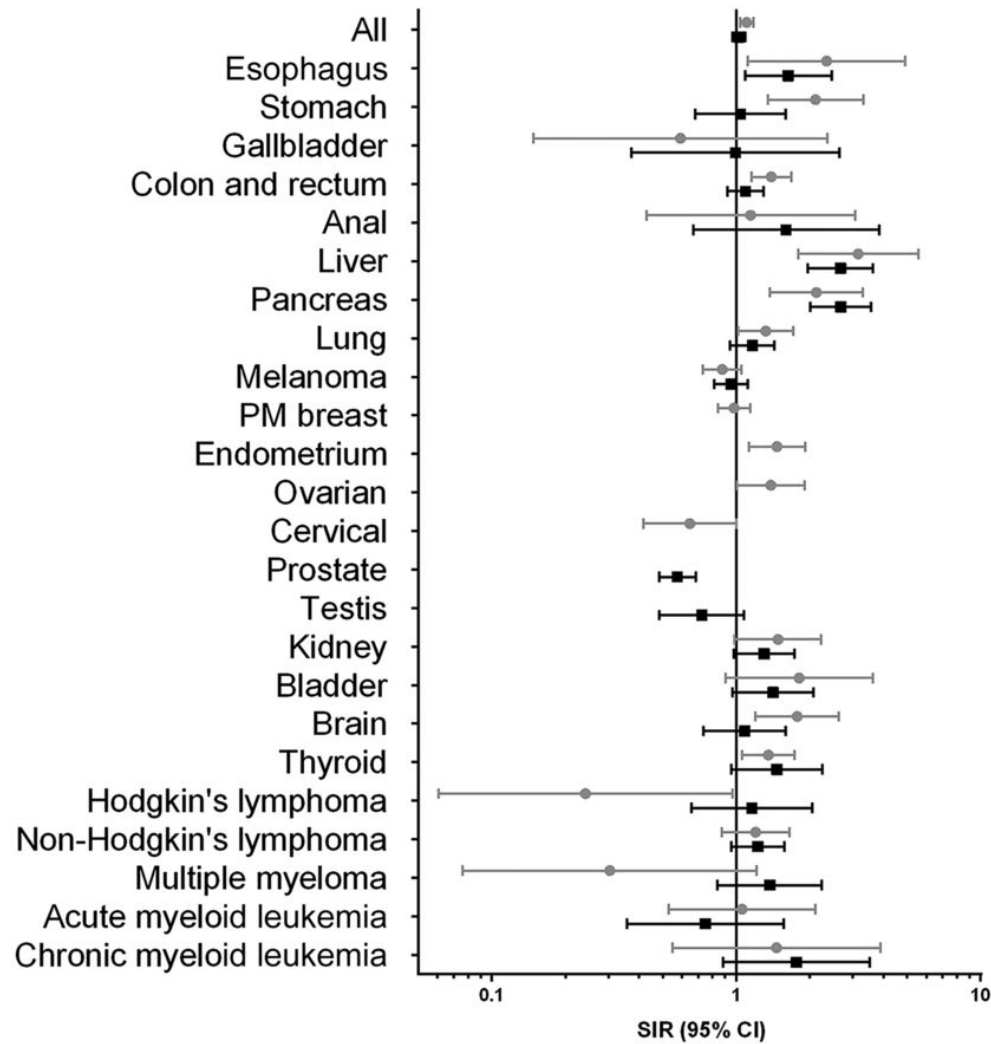
21 Neonatal encephalopathy

22 Malaria

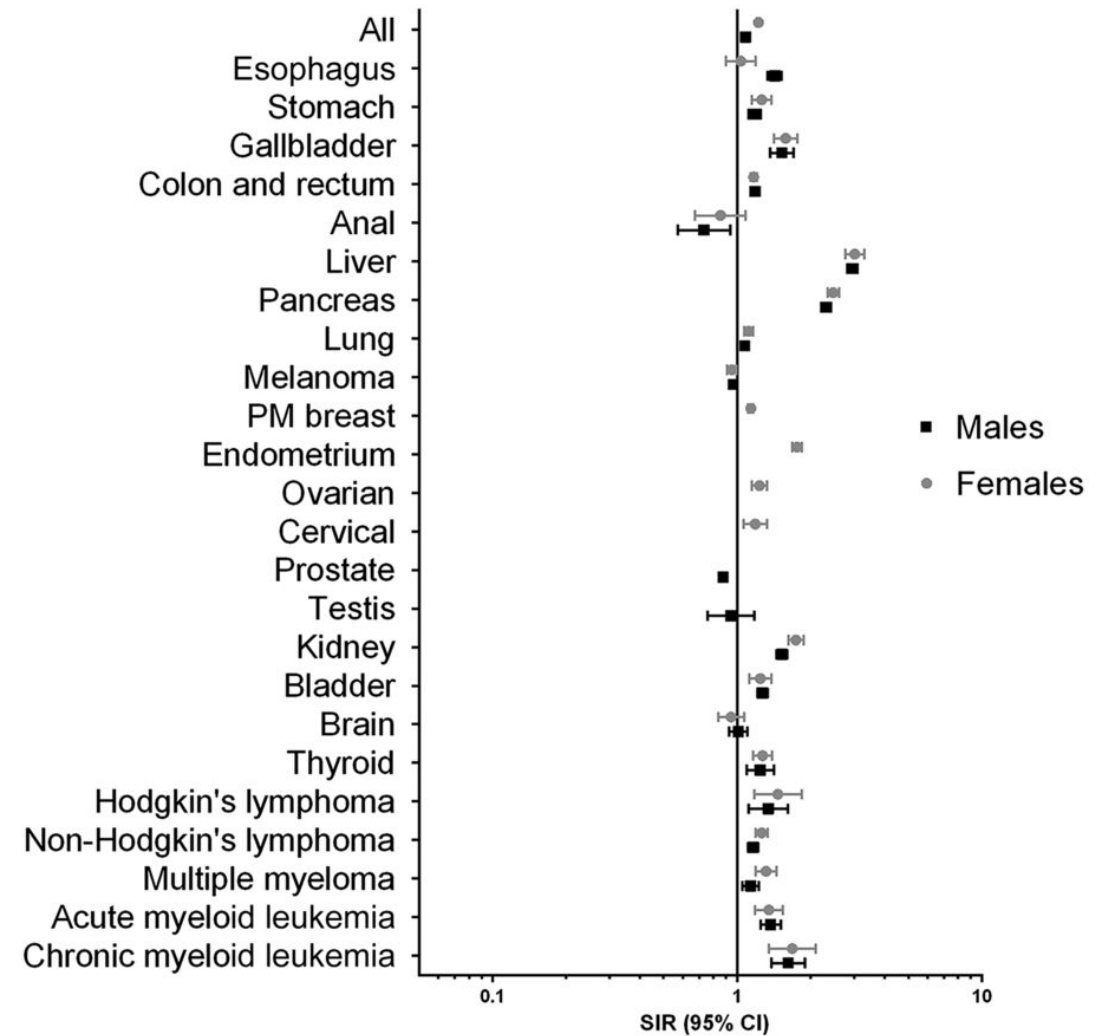
27 Neonatal sepsis

36 Other neonatal

A Type 1 Diabetes



B Type 2 Diabetes



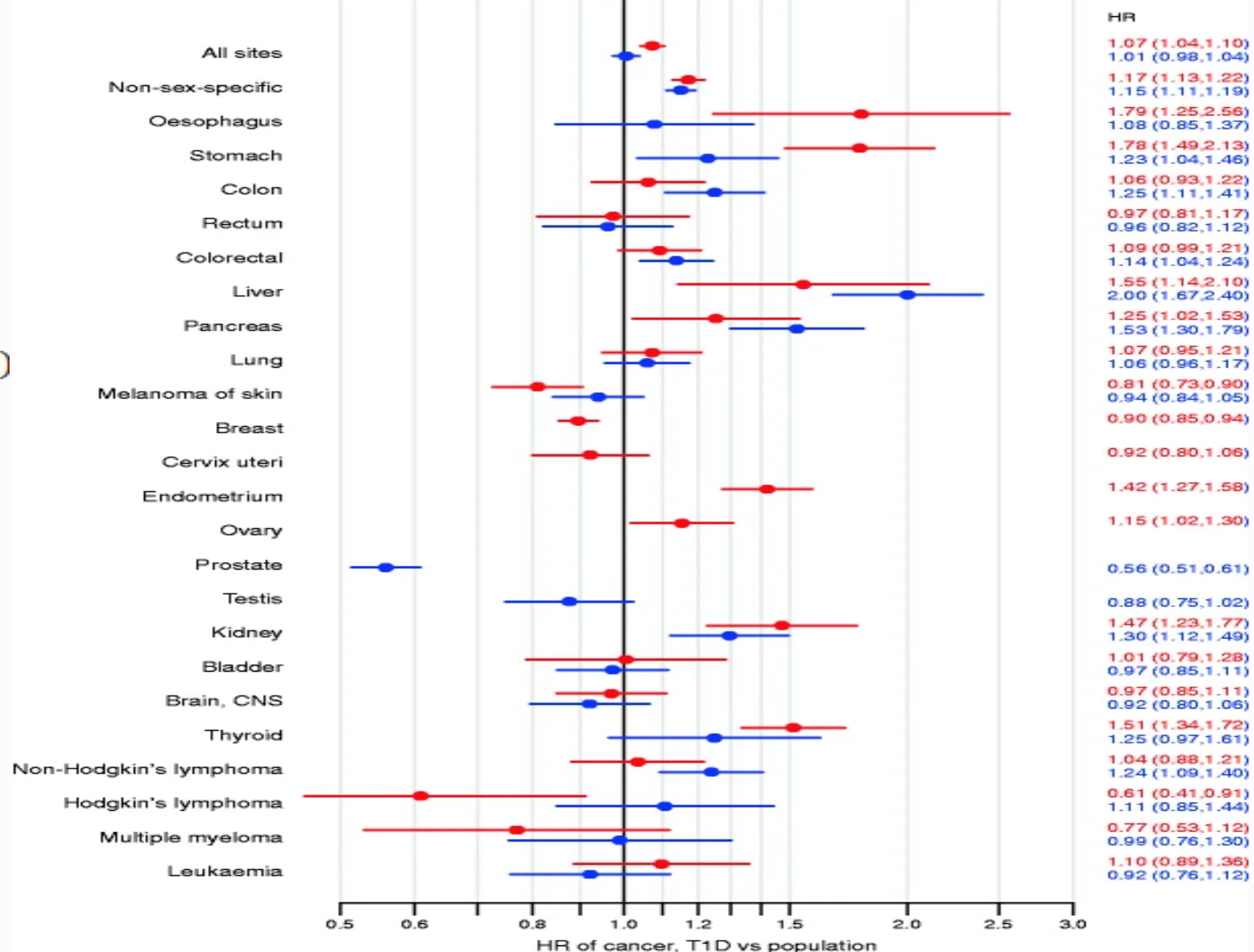
Cancer Risk Among People With Type 1 and Type 2 Diabetes: Disentangling True Associations, Detection Bias, and Reverse Causation

Jessica L. Harding; Jonathan E. Shaw; Anna Peeters; Bendix Cartensen; Dianna J. Magliano

Diabetes Care 2015;38(2):264–270

men (blue)

women (red)

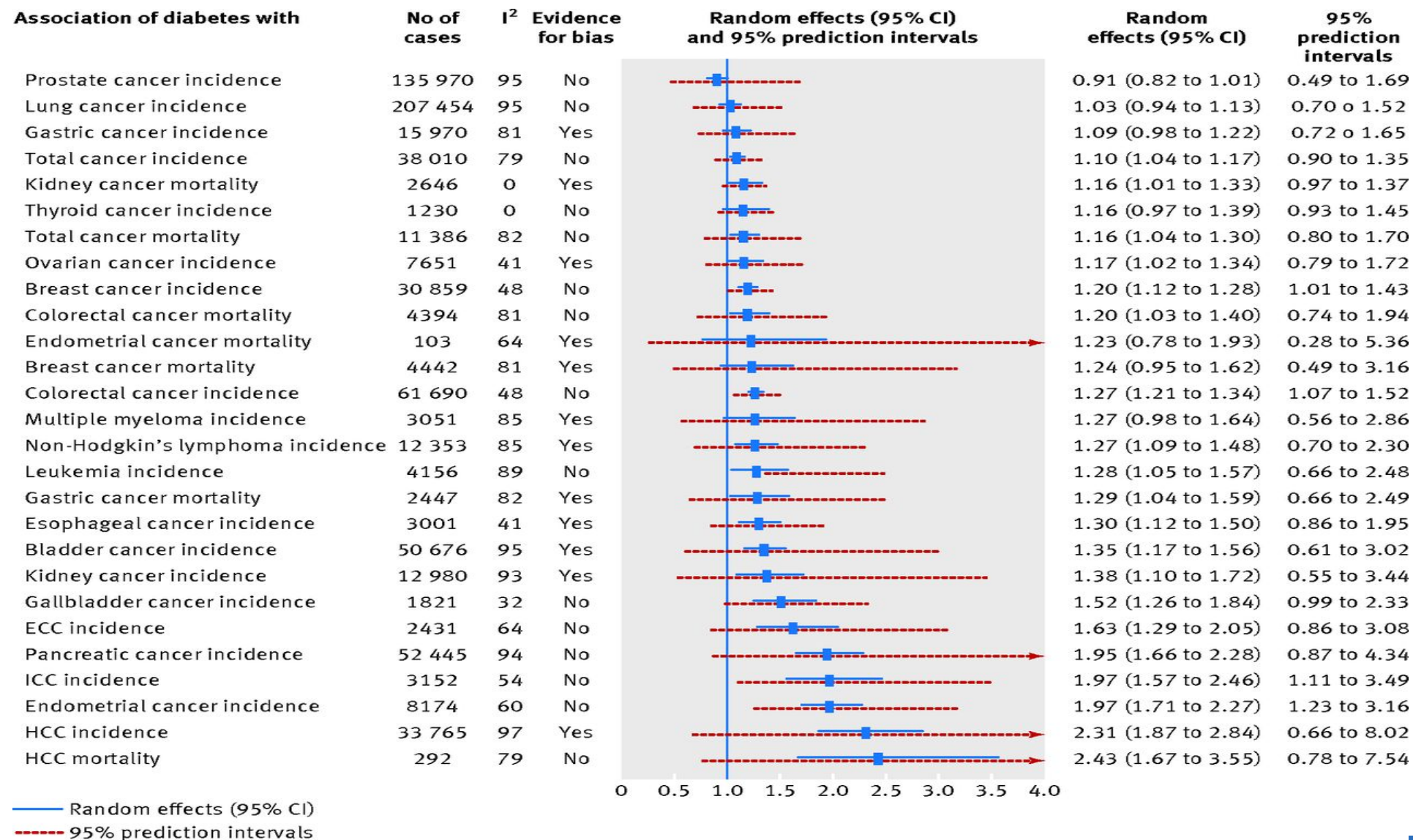


Carstensen, B.; Read, S.H.; Friis, S.; Sund, R.; Keskimäki, I.; Svensson, A.-M.; Ljung, R.; Wild, S.H.; Kerssens, J.J.; Harding, J.L.; et al.

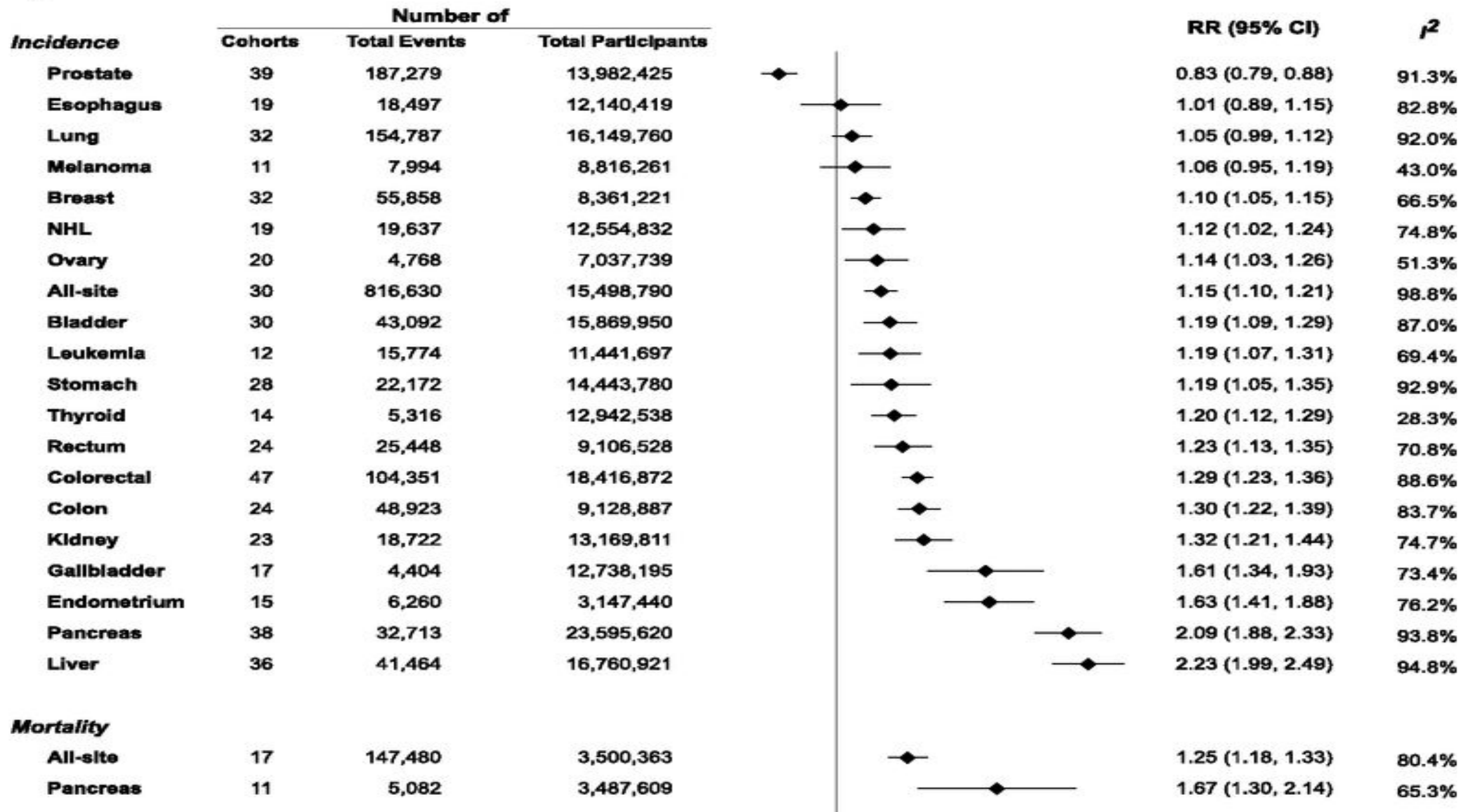
Diabetes and Cancer Research Consortium. Cancer incidence in persons with type 1 diabetes: A five-country study of 9000 cancers in type 1 diabetic individuals.

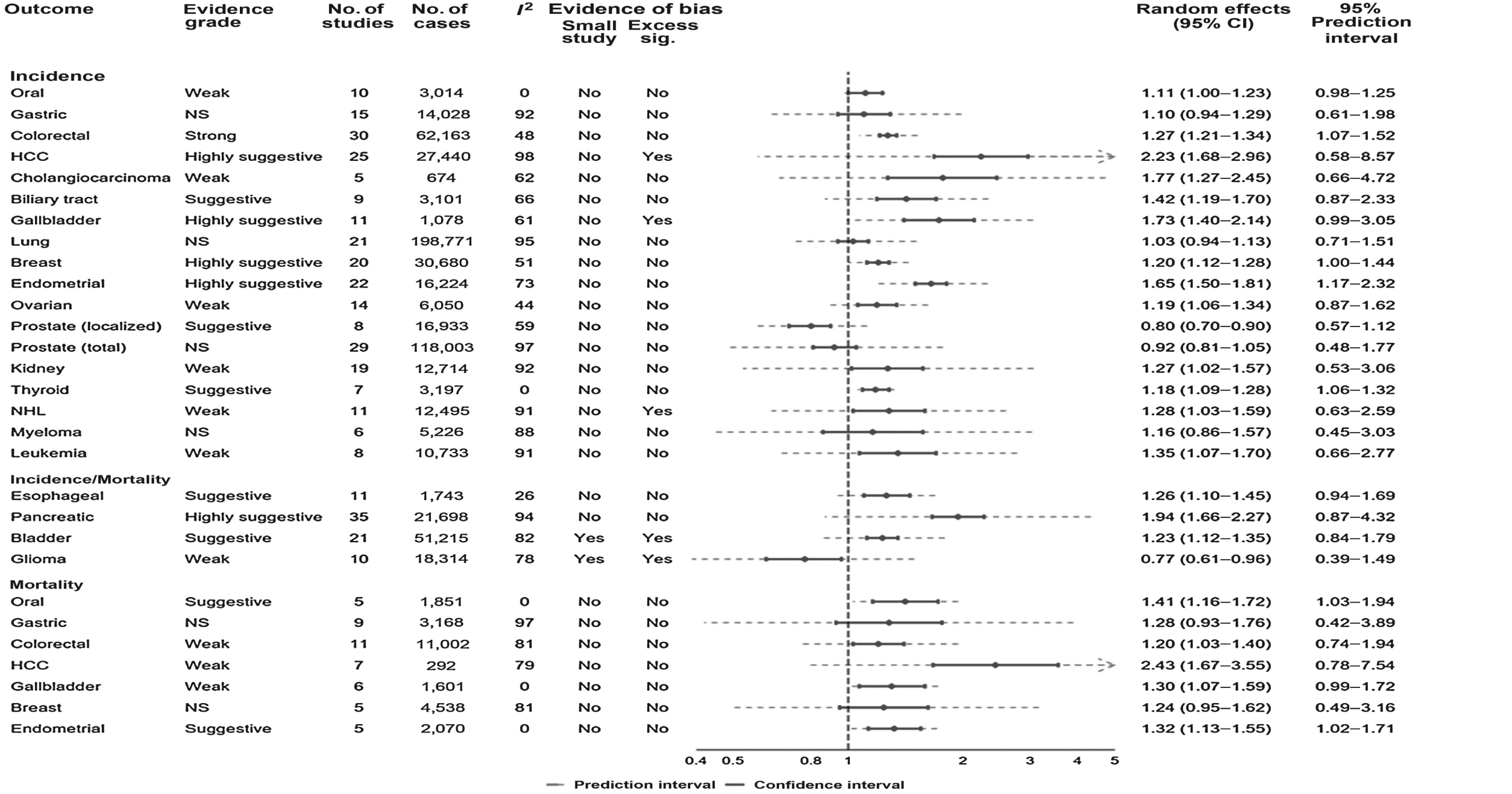
Diabetologia 2016, 59, 980–988.

Fig 2 Summary random effects estimates with 95% confidence and prediction intervals from 27 meta-analyses of type 2 diabetes and incidence of cancer or mortality (evidence for bias assessed with extrapolation of Egger regression line to study of infinite sample size and with excess significance test).



Konstantinos K Tsilidis et al. BMJ 2015;350:bmj.g7607





Type 2 Diabetes and Cancer: An Umbrella Review of Observational and Mendelian Randomization Studies

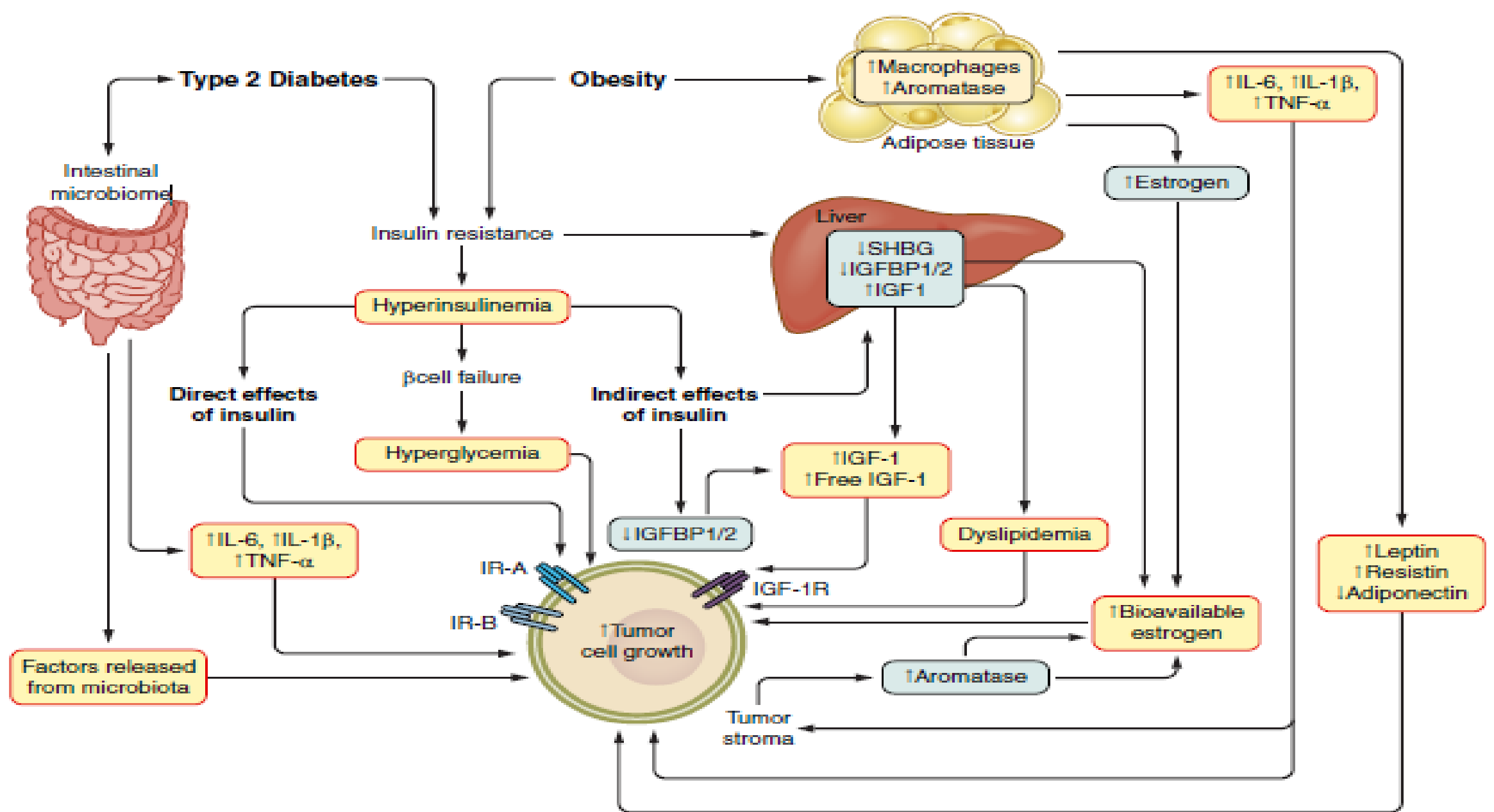
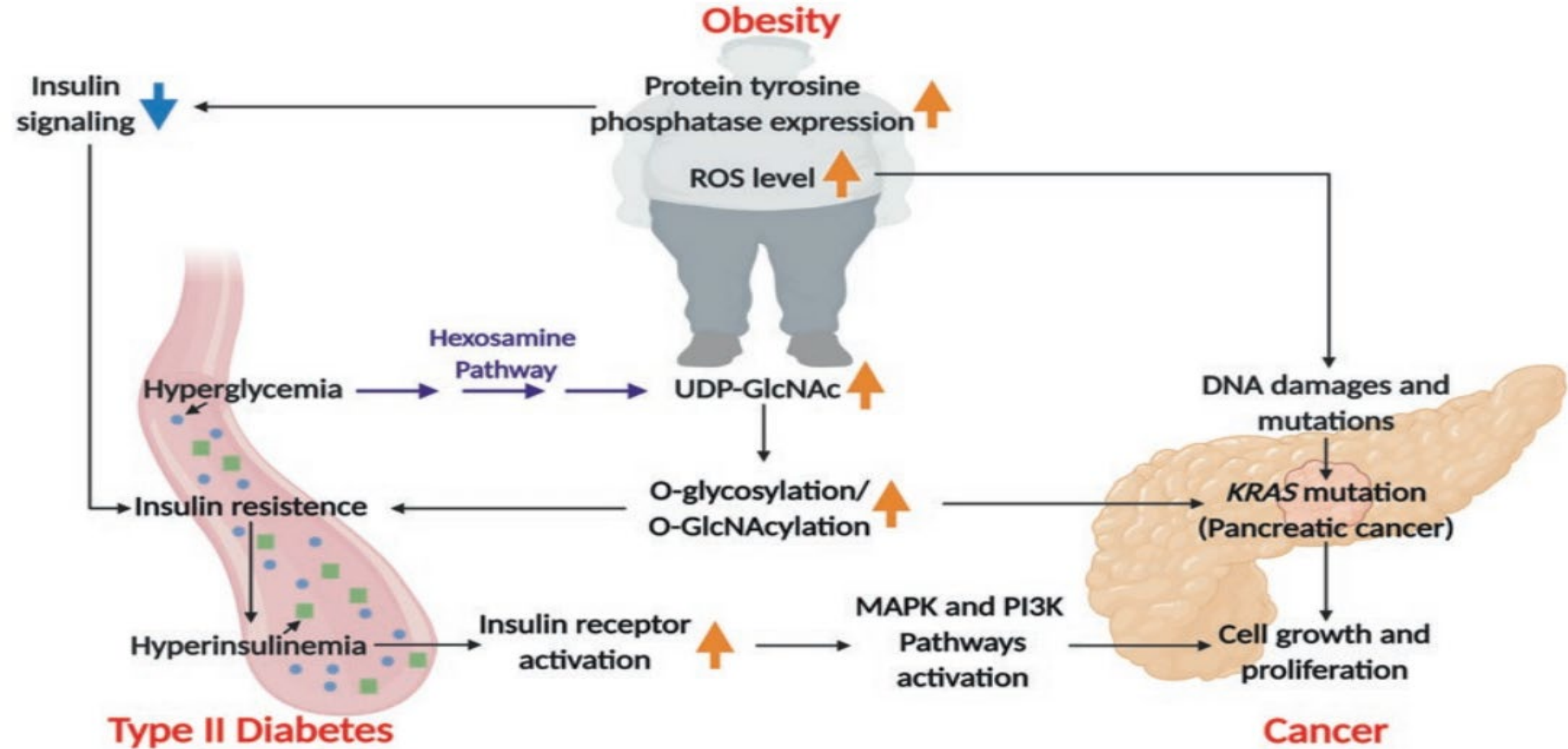
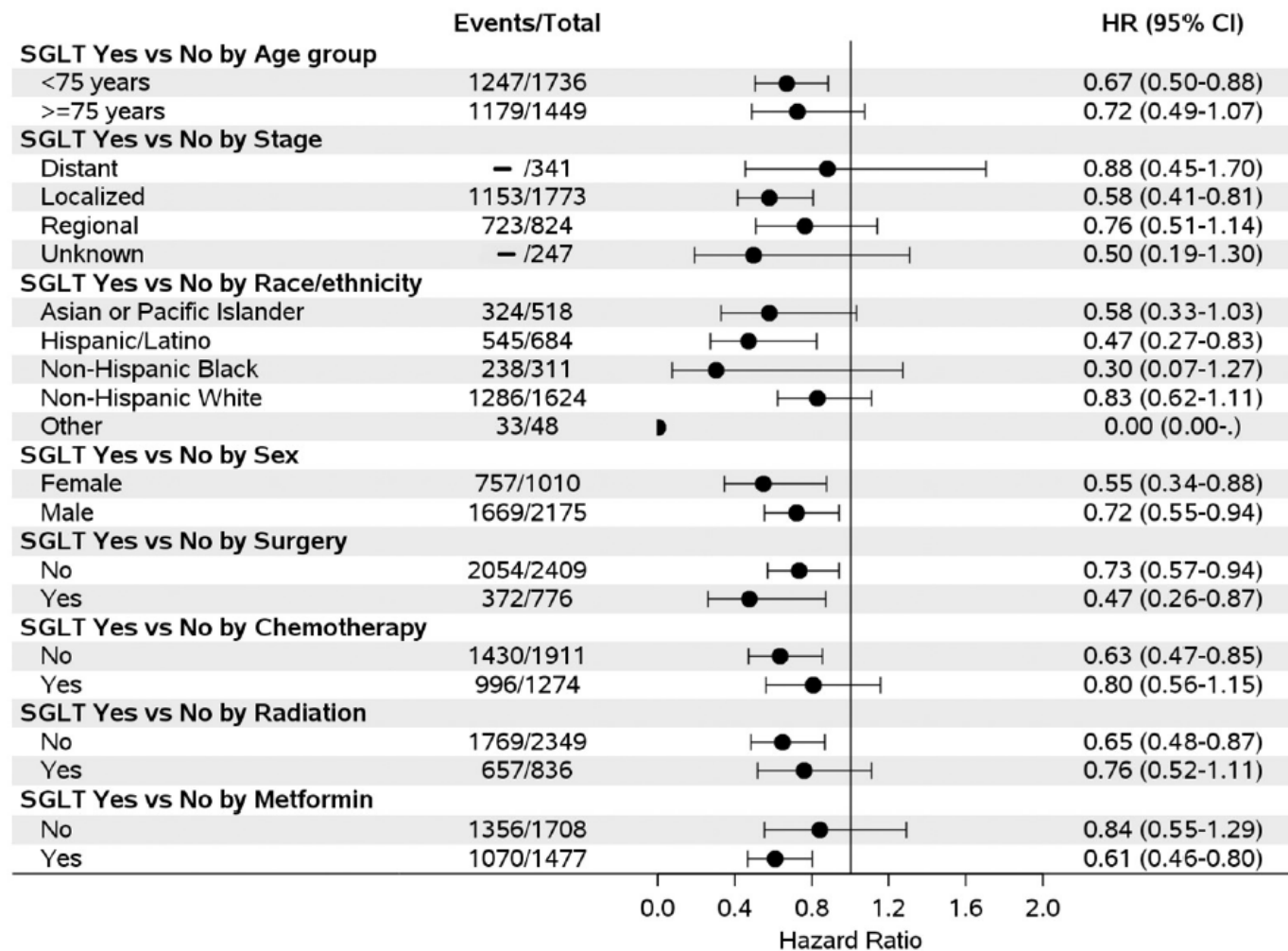


FIGURE 1. Mechanisms linking obesity, type 2 diabetes, and cancer.

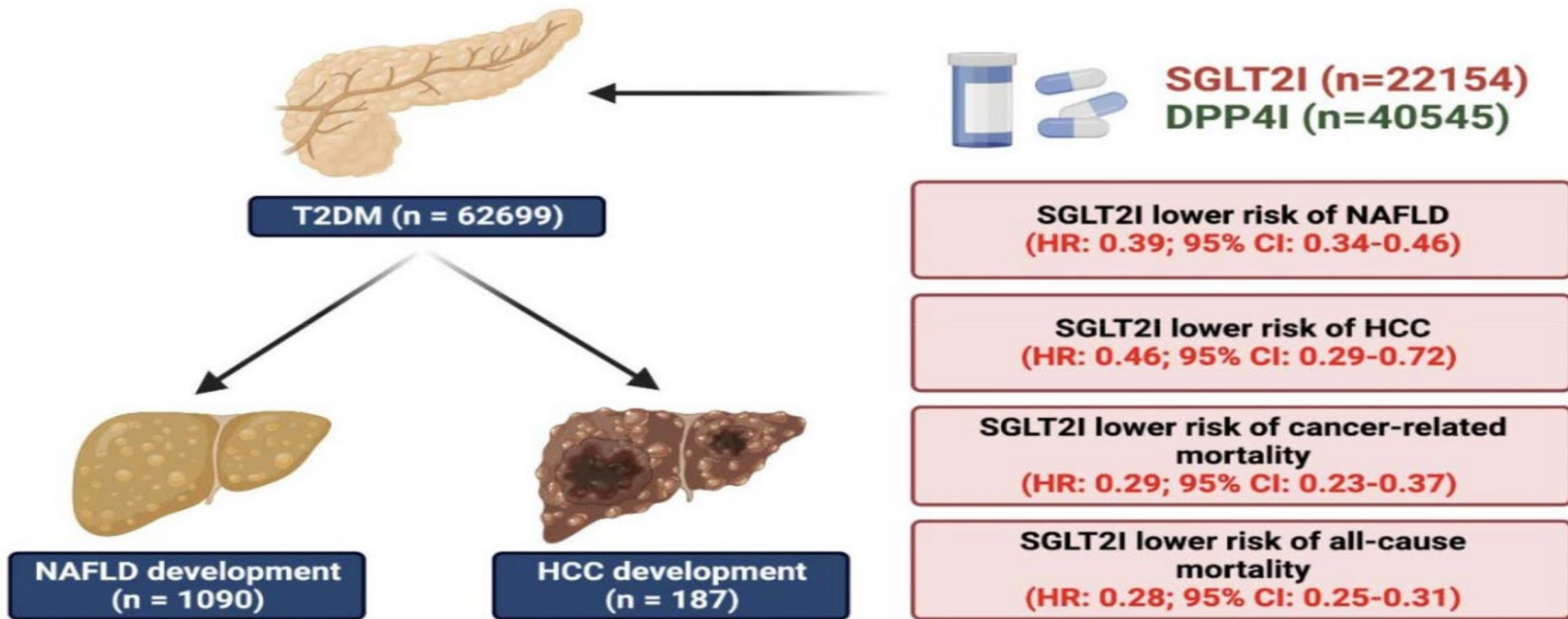


3185 pz con hcc 143 usavano SGLT2 (4,3 %)
 2426 morti no 2348 si 78

	Sensitivity analysis 1	
	Deaths/Patients	HR (95% CI)
SGLT2 inhibitor use		
No	86/137	Reference
Yes	78/137	0.50 (0.37–0.69)
Duration of use		
<12 months	51/85	0.52 (0.37–0.75)
≥ 12 months	27/52	0.47 (0.30–0.74)



Retrospective, territory-wide cohort study of T2DM patients treated with SGLT2i or DPP4i between 1st, 2015 January, and 31st December 2019

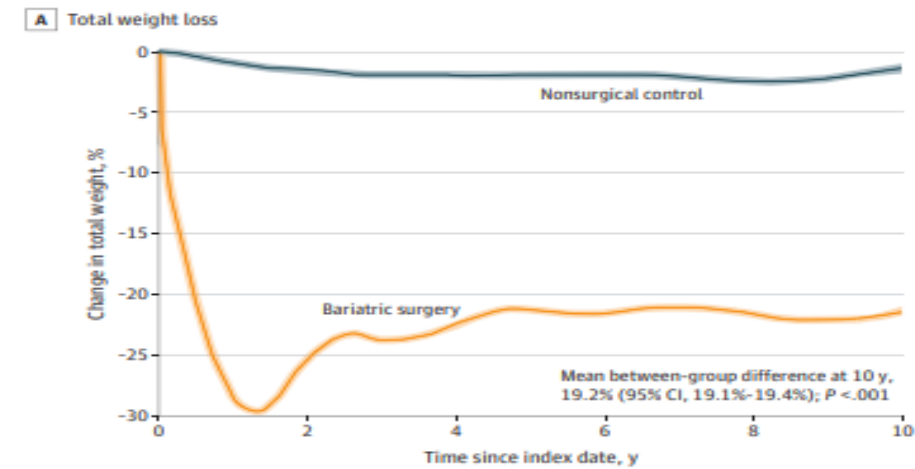


Patients on SGLT2i had a lower risk of NAFLD, and hepatocellular carcinoma compared to DPP4i after propensity scores matching and adjustments.

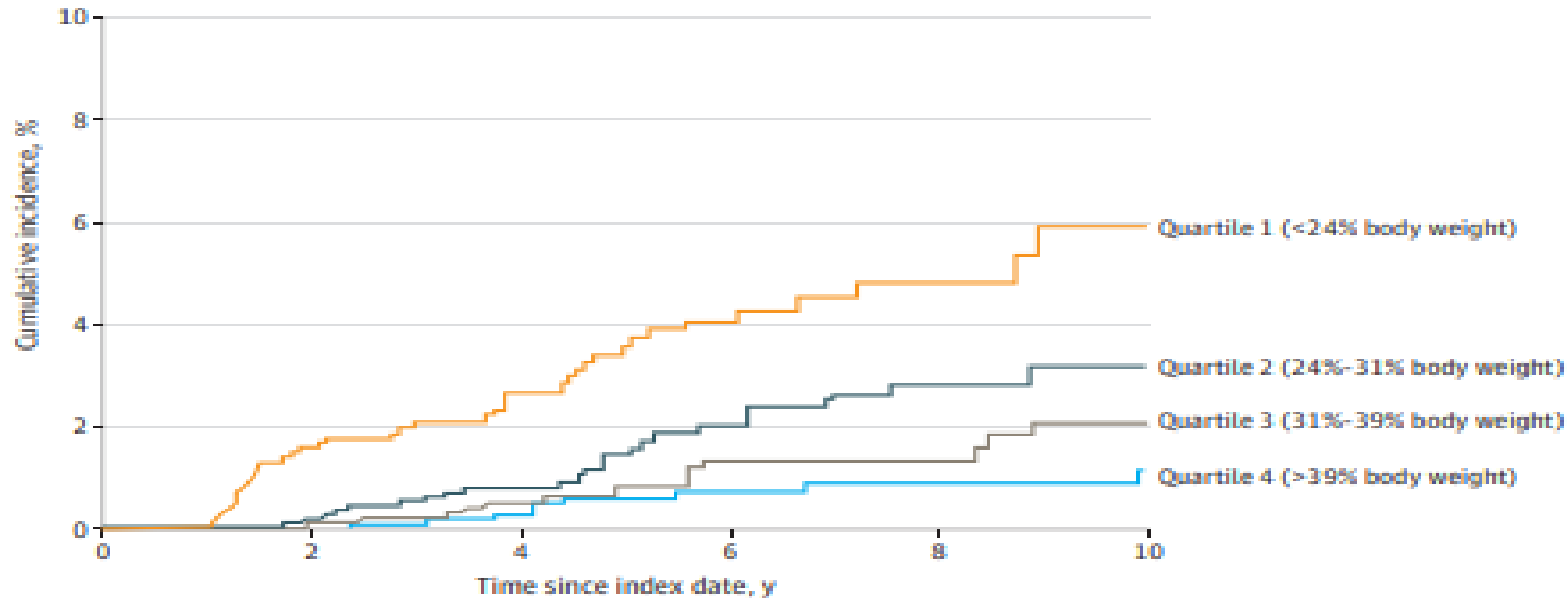
Lower risks of sodium glucose cotransporter 2 (SGLT2) inhibitors compared to dipeptidyl peptidase-4 (DPP4) inhibitors for new-onset non-alcoholic fatty liver disease and hepatocellular carcinoma in type 2 diabetes mellitus: A population-based study

Oscar Hou In Chou, Jing Ning, Raymond Ngai Chiu Chan, Cheuk To Chung, Helen Huang, Kenrick Ng, Edward Christopher Dee, Sharen Lee, Apichat Kaewdech, Tong Liu, Fengshi Jing, Bernard Man YungCheung, Gary Tse, Jiandong Zhou

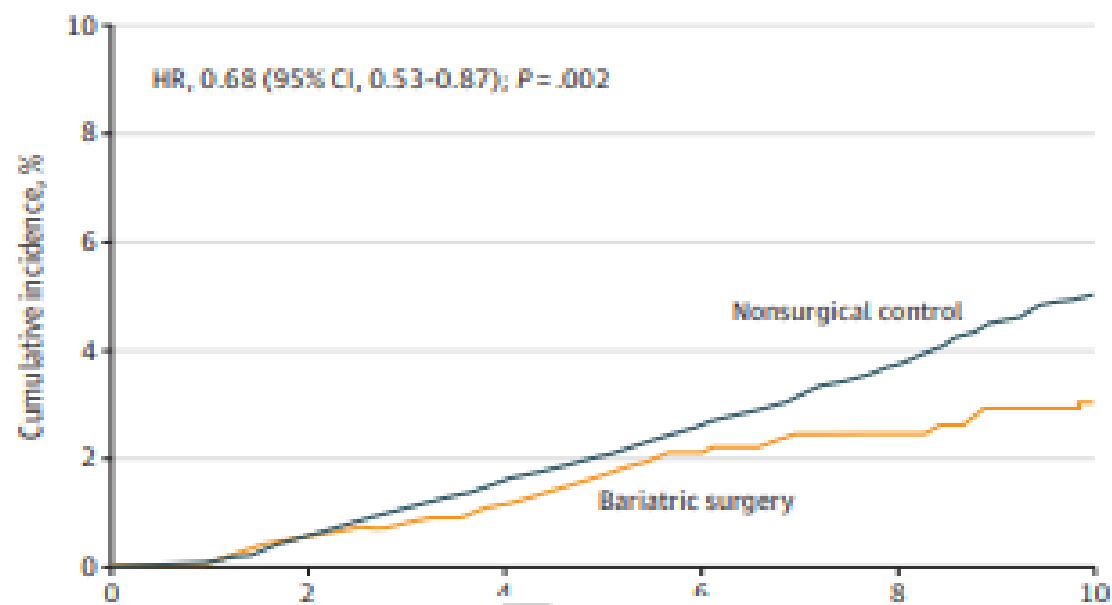
Figure 4. Weight Loss and Cumulative Incidence of Primary End Point Stratified by Maximum Weight Loss Quartile



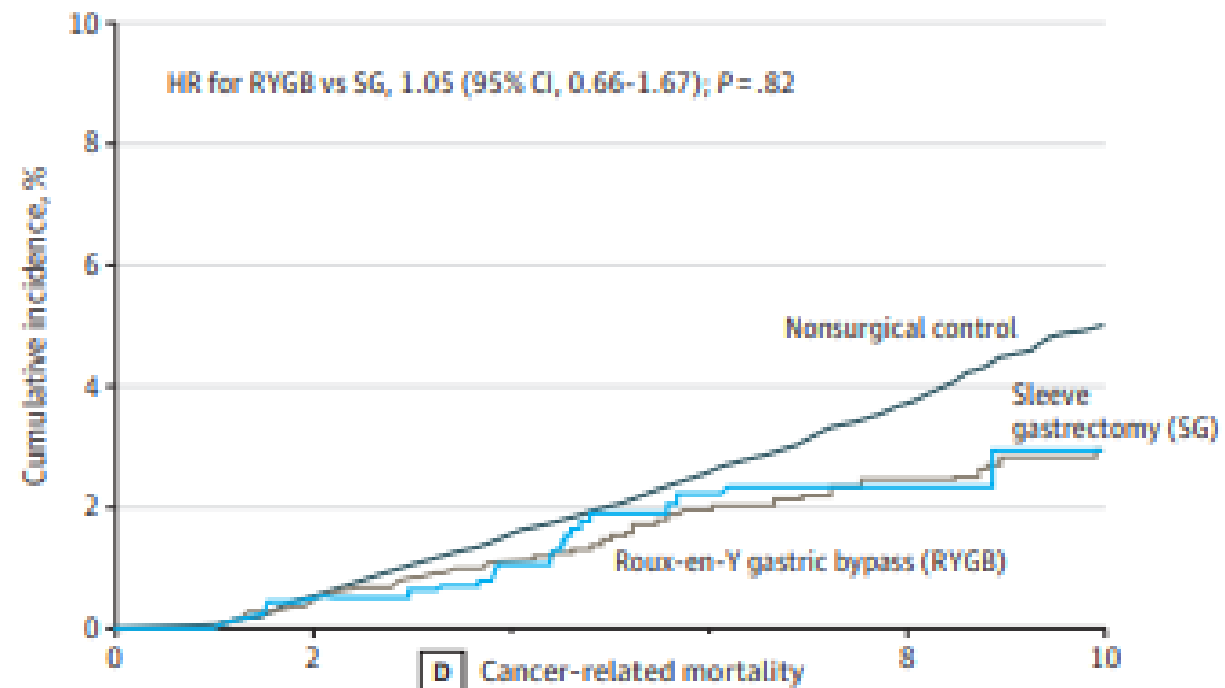
Obesity-associated cancer cases by surgically induced maximum weight loss quartiles



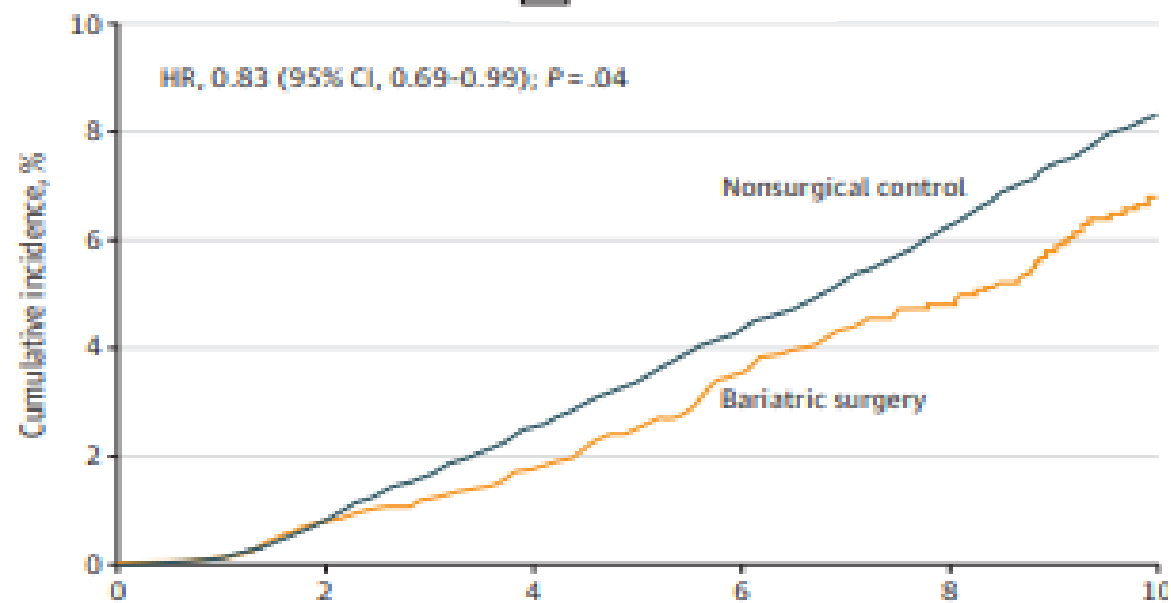
A Obesity-associated cancer cases



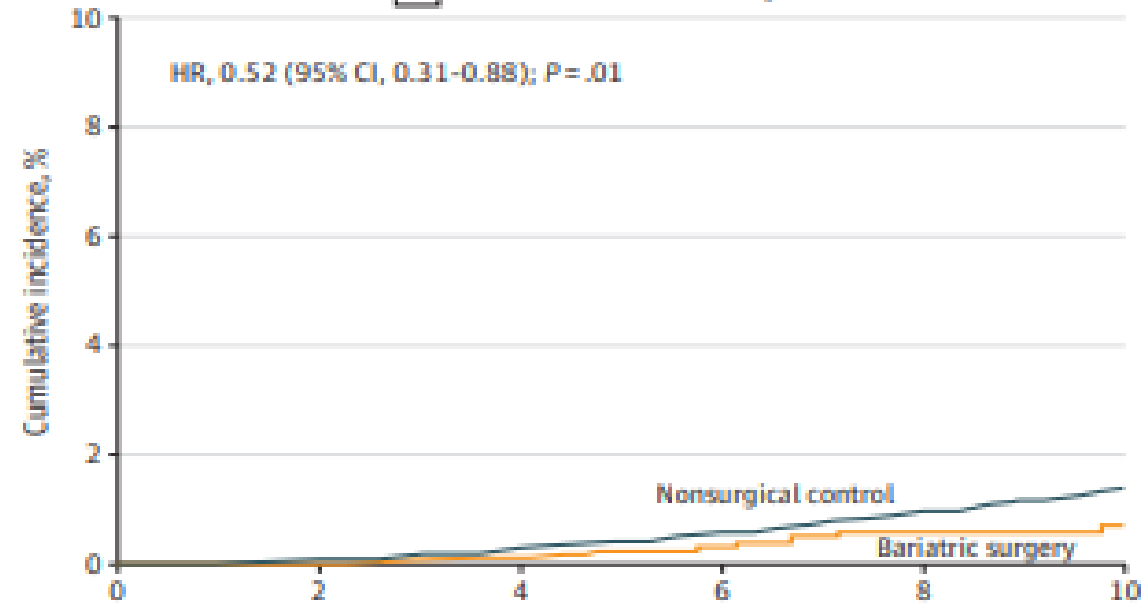
B Obesity-associated cancer cases by surgery type



C Total cancer cases



D Cancer-related mortality



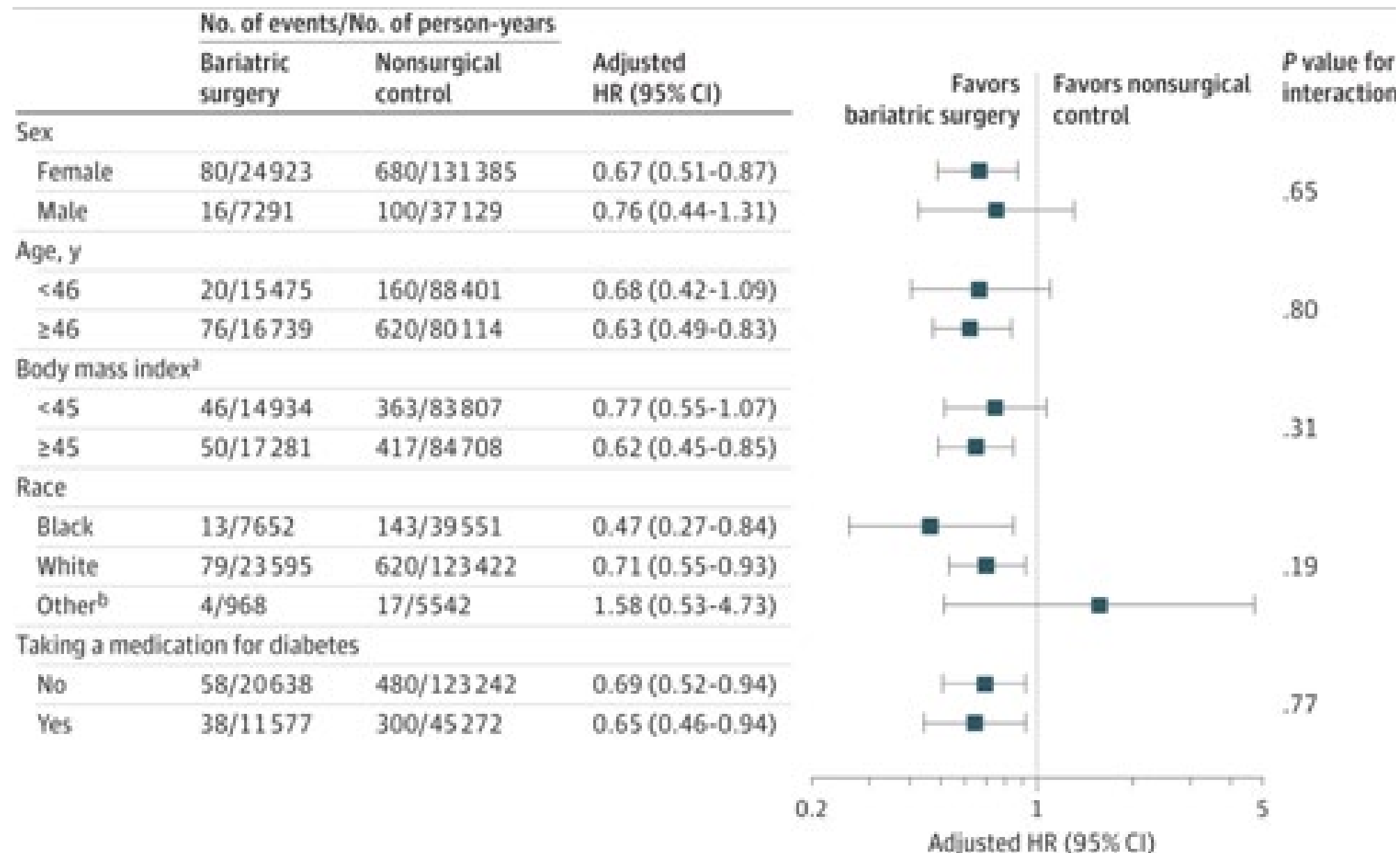


Tabella 1 ♦ **Effetti dei farmaci antitumorali sull'omeostasi glicemica**

FARMACI	GLICEMIA	HBA1C	MECCANISMO
Agenti ormonali - Agonisti GnRH (<i>triptorelina, leuprorelina</i>) - Anti-androgeni (<i>ciproterone acetate, flutamide, bicalutamide, nilutamide</i>) K prostata	↑ [64]	↑ [68]	Aumentata Ins-Res
Glucocorticoidi (<i>prednisone, prednisolone, metilprednisolone, desametasone</i>) K mammella, neuroendocrini, renali	↑ [70, 71]	↑ [58, 71]	Aumentata Ins-Res
Inibitori mTOR (<i>everolimus, sirolimus, temsirolimus</i>)	↑ [77]	↑ [58]	Aumentata Ins-Res
ICIs K polmonare, testa, collo, vescica, linfomi, melanoma - Anticorpi Anti-CTLA-4 (<i>ipilimumab</i>) - Inibitori PDL-1 (<i>atezolizumab, avelumab</i>) - Inibitori PD-1 (<i>nivolumab, pembrolizumab</i>)	↑ [89]	↑ [79]	Ridotta P/S Ins
Analoghi della somatostatina (<i>octreotide, lanreotide, pasireotide</i>) T neuroendocrini	↑ (pasireotide) [83] ↓ (octreotide) [84]	↑ [83]	Ridotta P/S Ins Ridotta P/S Gluc
L-asparaginasi Leucemia linfoblastica acuta	↑ [85, 86]		Ridotta P/S Ins
Agenti alchilanti (<i>cisplatino, carboplatino, oxaliplatino, ciclofosfamide, ecc.</i>) T. solidi	↑ [87-89]	↑ [58]	Ridotta P/S Ins?
Anti-microtubuli (<i>docetaxel, paclitaxel, vinorelbina, vincristina, ecc.</i>)	↑ [87, 90]	↑ [58]	Ridotta P/S Ins?
Anti-metaboliti (<i>5-fluorouracile, gemcitabina, capecitabina, metotrexate, ecc.</i>)	↑ [88, 91]	↑ [58]	Ridotta P/S Ins?
Antracicline (<i>doxorubicina, epirubicina, ecc.</i>)	↑ [87]		Ridotta P/S Ins?

Abbreviazioni: GnRH, gonadotropin-releasing hormone; Ins-Res, insulino-resistenza; HbA1c, emoglobina glicosilata; mTOR, mammalian target of rapamycin; ICIs, immune checkpoint inhibitors; CTLA-4, cytotoxic T-lymphocyte 4-antigen; PDL-1, programmed cell death ligand-1; PD-1 programmed cell death-1, P/S Ins, produzione/secrezione insulinica; P/S Gluc, produzione/secrezione glucaqone.

End-of-Life Care

Recommendations

13.19 When palliative care is needed in older adults with diabetes, providers should initiate conversations regarding the goals and intensity of care. Strict glucose and blood pressure control are not necessary **E**, and simplification of regimens can be considered. Similarly, the intensity of lipid management can be relaxed, and withdrawal of lipid-lowering therapy may be appropriate. **A**

13.20 Overall comfort, prevention of distressing symptoms, and preservation of quality of life and dignity are primary goals for diabetes management at the end of life. **C**

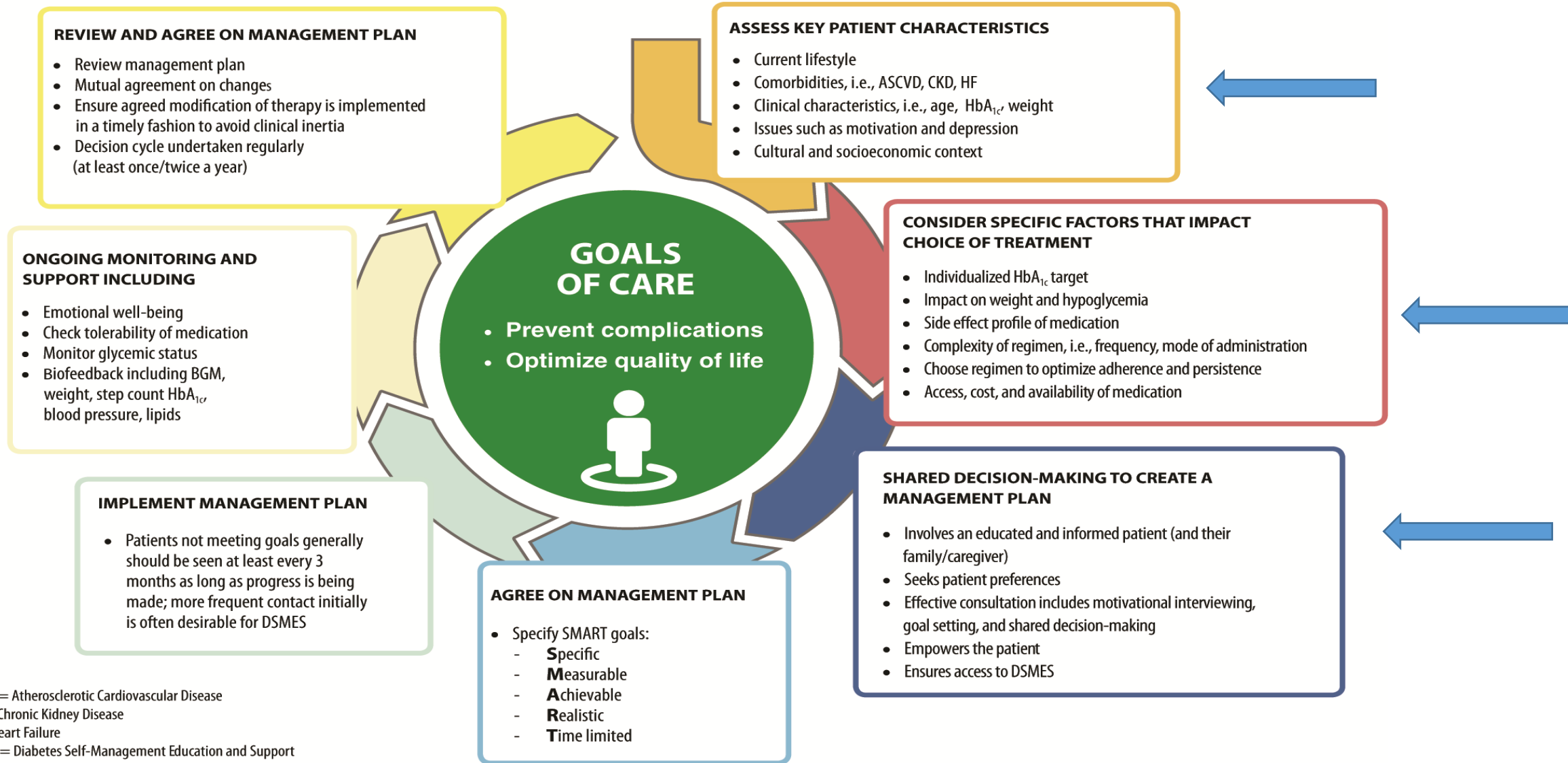
The management of the older adult at the end of life receiving palliative medicine or hospice care is a unique situation. Overall, palliative medicine promotes comfort, symptom control and prevention (pain, hypoglycemia, hyperglycemia, and dehydration), and preservation of dignity and quality of life in patients with limited life expectancy (**111,115**). In the setting of palliative care, providers should initiate conversations regarding the goals and intensity of diabetes care; strict glucose and blood pressure control may not be consistent with achieving comfort and quality of life. Avoidance of severe hypertension and hyperglycemia aligns with the goals of palliative care. In a multicenter trial, withdrawal of statins among patients in palliative care was found to improve quality of life (**116–118**). The evidence for the safety and efficacy of deintensification protocols in older adults is growing for both glucose and blood pressure control (**88,119**) and is clearly relevant for palliative care. A patient has the right to refuse testing and treatment, whereas providers may consider withdrawing treatment and limiting diagnostic testing, including a reduction in the frequency of blood glucose monitoring (**120,121**). Glucose targets should aim to prevent hypoglycemia and hyperglycemia. Treatment interventions need to be mindful of quality of life. Careful monitoring of oral intake is warranted. The decision process may need to involve the patient, family, and caregivers, leading to a care plan that is both convenient and effective for the goals of care (**122**). The pharmacologic therapy may include oral agents as first line, followed by a simplified insulin regimen. If needed, basal insulin can be implemented, accompanied by oral agents and without rapid-acting insulin. Agents that can cause gastrointestinal symptoms such as nausea or excess weight loss may not be good choices in this setting. As symptoms progress, some agents may be slowly tapered and discontinued. Different patient categories have been proposed for diabetes management in those with advanced disease (**55**).

A stable patient: Continue with the patient’s previous regimen, with a focus on 1) the prevention of hypoglycemia and 2) the management of hyperglycemia using blood glucose testing, keeping levels below the renal threshold of glucose, and hyperglycemia-mediated dehydration. There is no role for A1C monitoring.

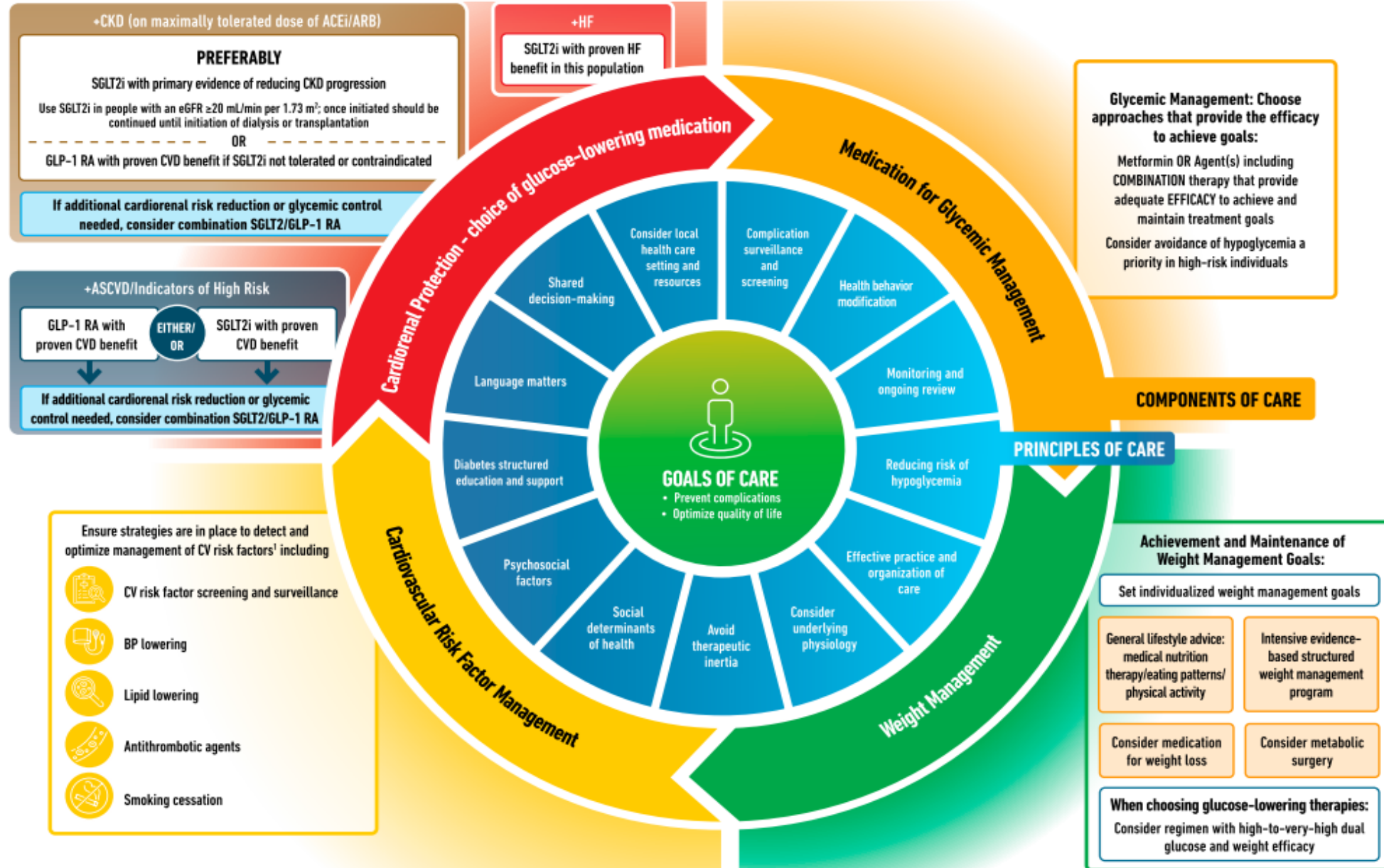
A patient with organ failure: Preventing hypoglycemia is of greatest significance. Dehydration must be prevented and treated. In people with type 1 diabetes, insulin administration may be reduced as the oral intake of food decreases but should not be stopped. For those with type 2 diabetes, agents that may cause hypoglycemia should be reduced in dose. The main goal is to avoid hypoglycemia, allowing for glucose values in the upper level of the desired target range.

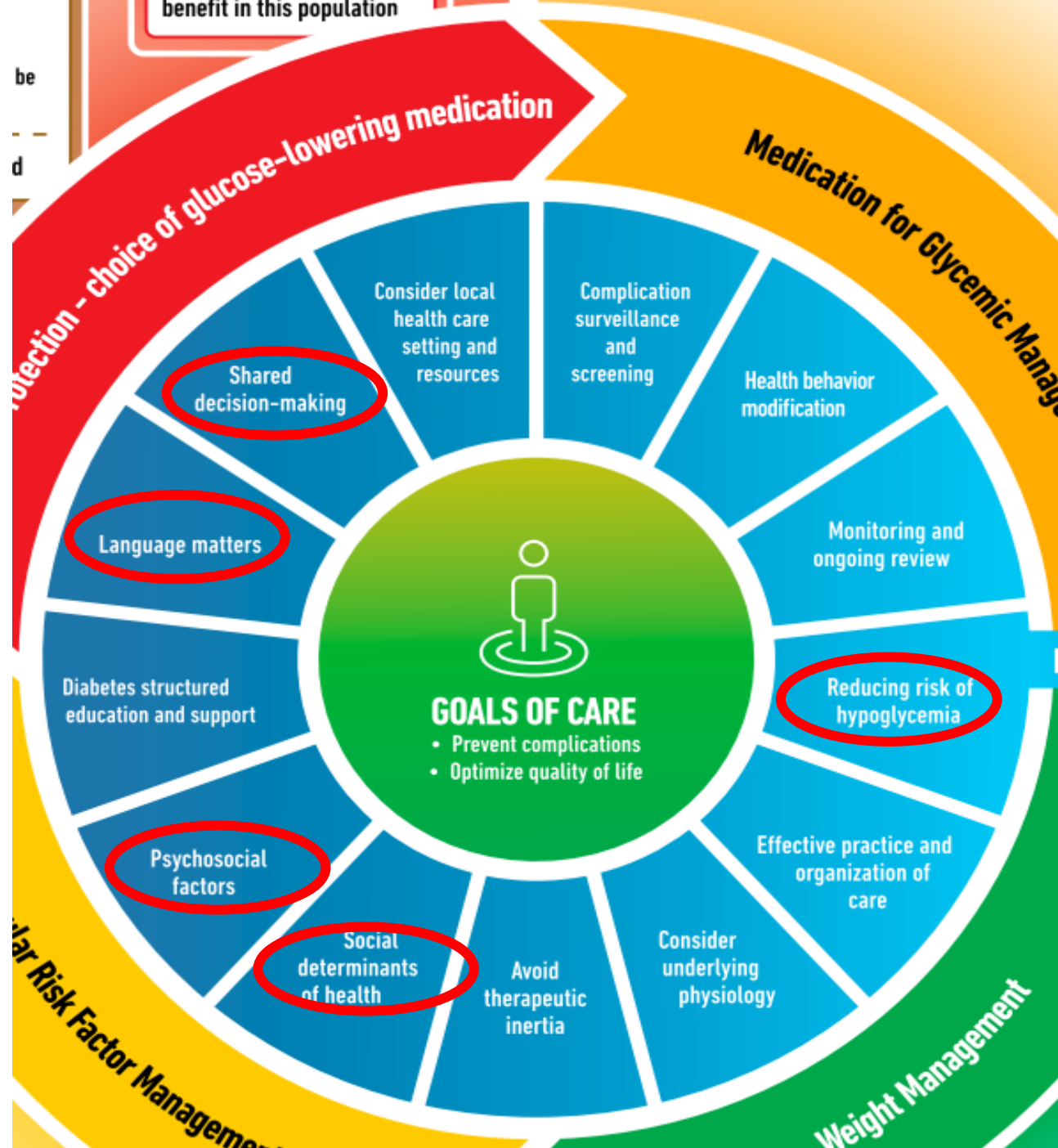
A dying patient: For patients with type 2 diabetes, the discontinuation of all medications may be a reasonable approach, as patients are unlikely to have any oral intake. In patients with type 1 diabetes, there is no consensus, but a small amount of basal insulin may maintain glucose levels and prevent acute hyperglycemic complications.

DECISION CYCLE FOR PATIENT-CENTERED GLYCEMIC MANAGEMENT IN TYPE 2 DIABETES

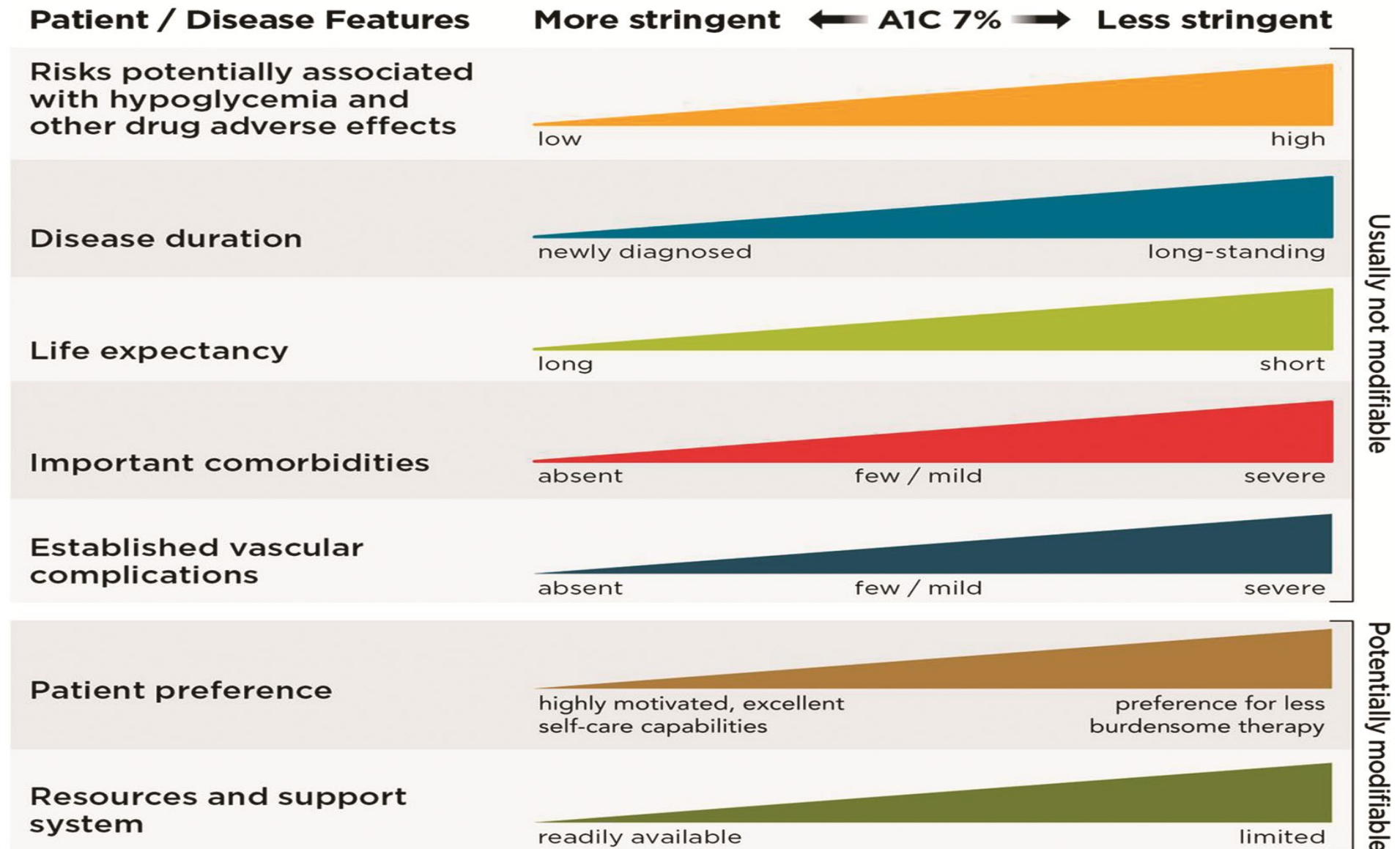


HOLISTIC PERSON-CENTERED APPROACH TO T2DM MANAGEMENT





Approach to Individualization of Glycemic Targets



Grado 1 Glicemia mattutina
< **160** mg/dl

Grado 2 Glicemia mattutina
> **160 mg/dl** < **250** mg/dl

Grado 3 Glicemia mattutina
> **250 mg/dl** < **500** mg/dl

Grado 4 Glicemia mattutina
> **500** mg/dl

Continuare trattamento

Dieta e modifiche sullo stile di vita

Trattamento con metformina se iperglicemia permane nonostante dieta e modifiche sullo stile di vita

Se sintomatico sospendere trattamento per 48-72 h fino al miglioramento

Dieta e modifiche sullo stile di vita

Trattamento con metformina

Se FBG > 200 dopo 2 settimane aggiungere un 2° ipoglicemizzante

Insulina per scarso controllo glicemico

Sospendere trattamento programmato e riprendere con glicemia < 250 e asintomatico

Modificare dose terapia

Idratazione intravenosa

Trattamento con metformina + un 2° agente ipoglicemizzante

Dieta e modifiche sullo stile di vita

Insulina per scarso controllo glicemico

Sospendere trattamento fino a risoluzione dei sintomi e riprendere trattamento con dose più bassa quando glicemia < 250

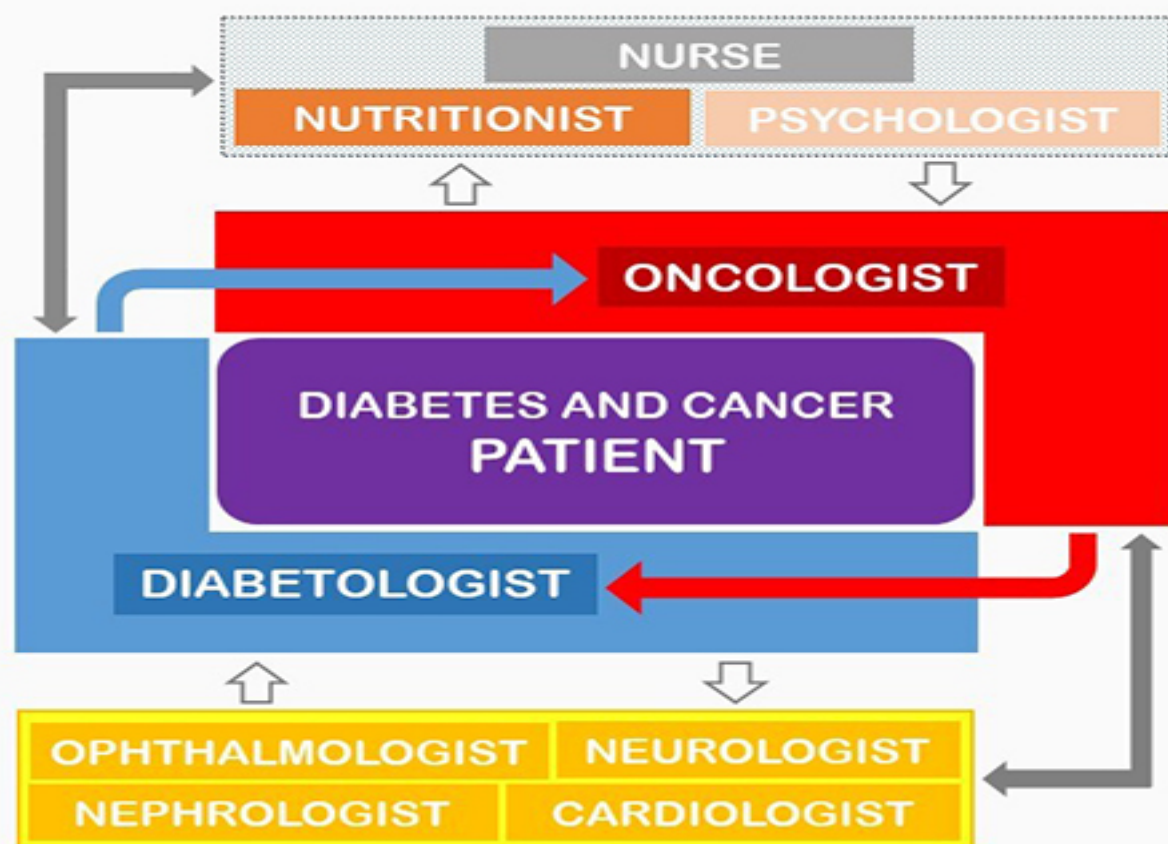
Idratazione intravenosa

Insulina per rapido controllo glicemico

Trattamento con metformina + un 2° agente ipoglicemizzante

Dieta e modifiche sullo stile di vita

ONCOMETABOLIC TEAM



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Diabete e cancro

Domenico Voce
Diabetologia Crotone

Grazie per l'attenzione



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