







# The 5<sup>th</sup> SOMS Spring Congress & AOASO Joint Symposium Incheon, 25 March 2023

# Management of Patients with DIABESITY:

Sharing Experiences from Indonesia

Dicky L. Tahapary

Division of Endocrinology, Metabolism and Diabetes, Department of Internal Medicine, Dr. Cipto Mangunkusumo National General Hospital, Faculty of Medicine Universitas Indonesia (RSCM-FKUI) Metabolic, Cardiovascular and Aging Cluster, The Indonesian Medical Education and Research Institute, Faculty of Medicine Universitas Indonesia (IMERI-FKUI)

# What is Diabesity?





# Diabesity describes the combined detrimental health effects of obesity and diabetes mellitus.

# **Dual Problem of Obesity and Type 2 Diabetes Mellitus**





The worldwide dual epidemic of obesity and type 2 diabetes is an important public health issue.

# **Problem of Obesity**





The estimates for **global levels of overweight and obesity (BMI ≥25kg/m2)**, also referred to as high BMI, suggest that **over 4 billion people may be affected by 2035**, compared with **over 2.6 billion in 2020**.

Numbers of people (aged over 5 years) and percentage of the population with overweight or obesity\*

	2020	2025	2030	2035
Number with overweight or obesity (BMI≥25kg/m²) (millions)	2,603	3,041	3,507	4,005
Number with obesity (BMI ≥30kg/m²) (millions)	988	1,249	1,556	1,914
Proportion of the population with overweight or obesity (BMI ≥25kg/m²)	38%	42%	46%	51%
Proportion of the population with obesity (BMI ≥30kg/m²)	14%	17%	20%	24%

<sup>\*</sup> For children and adolescents, overweight and obesity are defined using the WHO classification of +1SD and +2SD above median growth reference.

World Obesity Atlas 2023

# **Problem of Obesity**







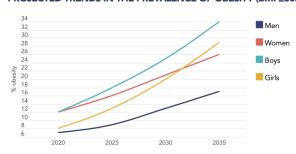
## Indonesia

ADULTS WITH OBESITY 2035

22%

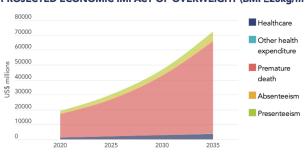
HIGH

#### PROJECTED TRENDS IN THE PREVALENCE OF OBESITY (BMI ≥30kg/m²)





#### PROJECTED ECONOMIC IMPACT OF OVERWEIGHT (BMI ≥25kg/m²)





OVERWEIGHT IMPACT ON NATIONAL GDP 2035 3.1% VERY HIGH

#### IMPACT OF OVERWEIGHT (BMI ≥25kg/m²) 2020-2035

	Healthcare impact of BMI ≥25kg/m², US\$ million	Total economic impact of BMI ≥25kg/m², US\$ million	Estimated GDP US\$ billion	Impact of BMI ≥25kg/m² on GDP
2020	1,265	18,950	1,096	1.7%
2025	1,884	29,854	1,416	2.1%
2030	2,724	46,761	1,826	2.6%
2035	3,837	72,178	2,355	3.1%



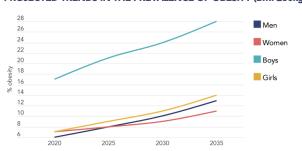
### **South Korea**

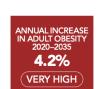
ADULTS WITH OBESITY 2035

12%

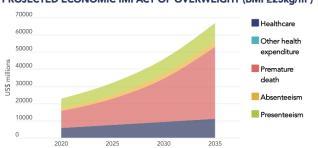
MEDIUM

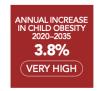
#### PROJECTED TRENDS IN THE PREVALENCE OF OBESITY (BMI ≥30kg/m²)





#### PROJECTED ECONOMIC IMPACT OF OVERWEIGHT (BMI ≥25kg/m²)





#### OVERWEIGHT IMPACT ON NATIONAL GDP 2035 2.2% VERY HIGH

#### IMPACT OF OVERWEIGHT (BMI ≥25kg/m²) 2020-2035

	Healthcare impact of BMI ≥25kg/m², US\$ million	Total economic impact of BMI ≥25kg/m², US\$ million	Estimated GDP US\$ billion	Impact of BMI ≥25kg/m² on GDP
2020	5,855	22,614	1,637	1.4%
2025	7,406	31,902	1,890	1.7%
2030	9,124	45,998	2,331	2.0%
2035	11,005	66,857	3,065	2.2%



World Obesity Atlas 2023

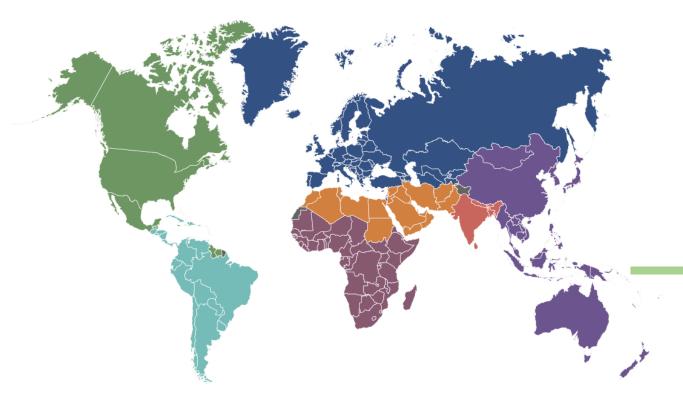
2045 49 million

2021 32 million









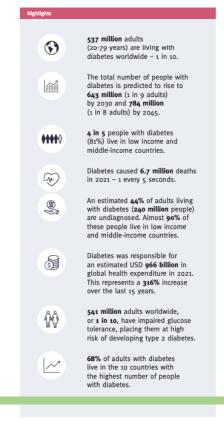
iddle East & North Africa (MENA)

2045 136 million

uth-East Asia (SEA)

2045 152 million

2030 113 million





The IDF Diabetes Atlas 10th edition has been produced thanks to educational grants from Novo Nordisk, Pfizer-MSD Alliance and Sanofi:







Need more information:



68%

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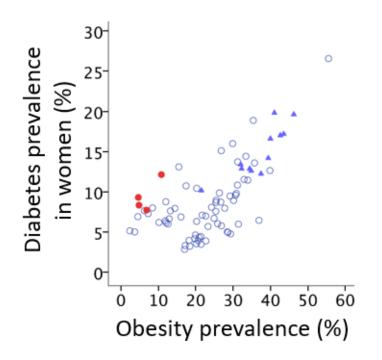


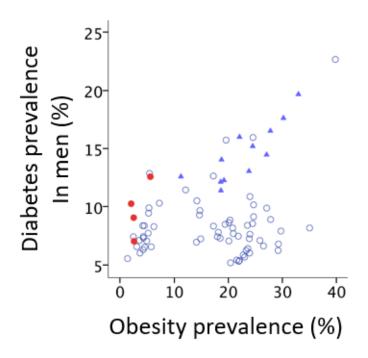
The prevalence of diabetes in Indonesia has almost doubled from 6.9% to 10.5% -- in parallel with the increased prevalence of obesity from ~20% to ~30%

# **Obesity and Type 2 Diabetes Mellitus**



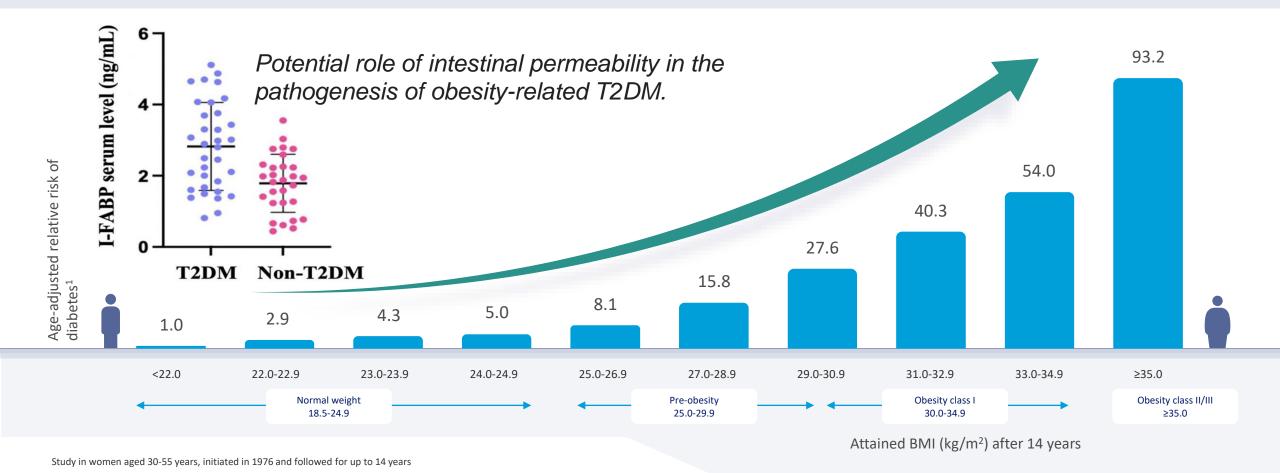
Increased adiposity is the strongest risk factor for developing diabetes.





# The relative risk for developing T2D increases with increasing body mass index





Colditz et al. Ann Intern Med. 1995 Apr 1;122(7):481-6.

PLoS ONE 2023 18(1): e0279915

# **DIABESITY "Diabetes and Obesity" in Indonesia**









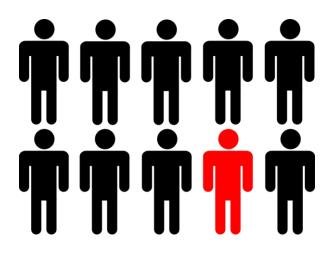
~70%

people with diabetes have overweight or obesity in Indonesia

## **Diabetes Mellitus in Indonesia**







**Diabetes Mellitus** 

### ~ 20 Million People With Diabetes in Indonesia

~86% undiagnosed ~70% uncontrolled ~ 25% less than 45 years old

### High proportion of diabetes complications

~20% cardiovascular ~40% diabetic kidney disease ~ 30% eye complications



# **Obesity & Other Cardiometabolic Risk Factors in Indonesia**









Obesity

Prediabetes
Dyslipidemia
Hypertension

Metabolic Syndrome

# Cardiometabolic-Based Chronic Disease (CMBCD)





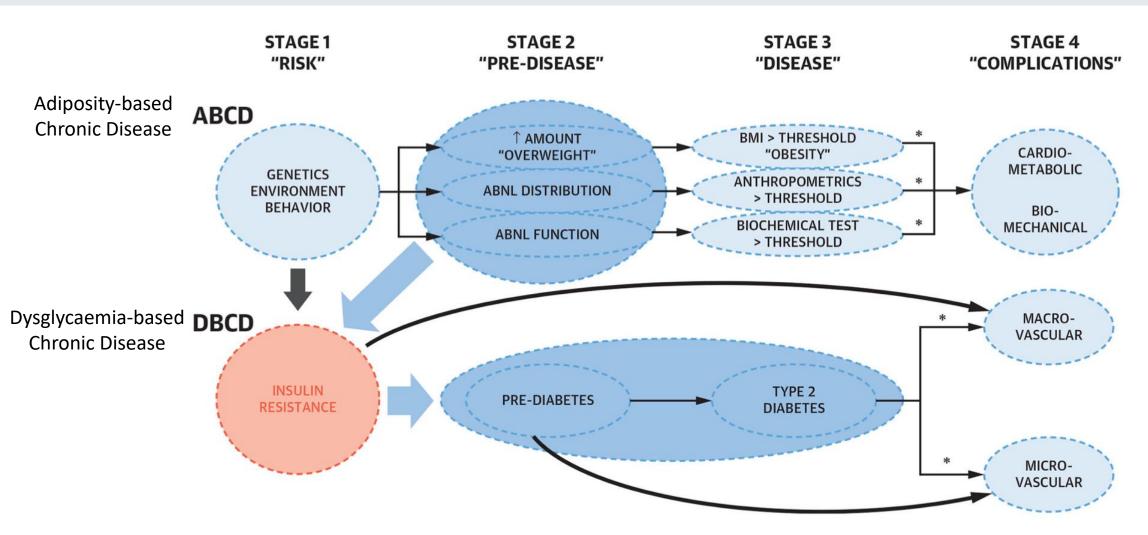
	CARDIOMETABOLIC-BASED CHRONIC DISEASE STAGE 1 "RISK"	CARDIOMETABOLIC-BASED CHRONIC DISEASE STAGE 2 "PRE-DISEASE"	CARDIOMETABOLIC-BASED CHRONIC DISEASE STAGE 3 "DISEASE" EARLY NO SYMPTOMS	CARDIOMETABOLIC-BASED CHRONIC DISEASE STAGE 4 "COMPLICATIONS" LATE SYMPTOMS
PRIMARY DRIVERS	GENETICS ENVIRONMENT BEHAVIOR	PRE-DISEASE PHENOTYPE	DISEASE PHENOTYPE	COMPLICATION PHENOTYPE
METABOLIC DRIVER 1: ADIPOSITY-BASED CHRONIC DISEASE	ADIPOSITY AMOUNT ADIPOSITY DISTRIBUTION ADIPOSITY FUNCTION	OVERWEIGHT HIGH WAIST CIRCUMFERENCE ABNORMAL BIOMARKERS	OBESITY	METABOLIC BIOMECHANICAL
METABOLIC DRIVER 2: DYSGLYCEMIA-BASED CHRONIC DISEASE	INSULIN RESISTANCE β-CELL DYSFUNCTION	PRE-DIABETES	TYPE 2 DIABETES	MICROVASCULAR MACROVASCULAR
CARDIOVASCULAR DISEASE	HYPERTENSION DYSLIPIDEMIA	SUBCLINICAL CHD LV DYSFUNCTION LA ABNORMALITY	ASYMPTOMATIC CHD ASYMPTOMATIC HF ASYMPTOMATIC AF	SYMPTOMATIC CHD SYMPTOMATIC HF SYMPTOMATIC AF

J Am Coll Cardiol. 2020 Feb 11;75(5):525-538.

# Cardiometabolic-Based Chronic Disease (CMBCD)







### EOSS: EDMONTON OBESITY STAGING SYSTEM - Staging Tool





#### STAGE 0

- · NO sign of obesity-related risk factors
- · NO physical symptoms
- NO psychological symptoms
- NO functional limitations

#### Case Example:

Physically active female with a BMI of 32 kg/m², no risk factors, no physical symptoms, no self-esteem issues, and no functional limitations.



#### STAGE 1

- Patient has obesity-related SUBCLINICAL risk factors (borderline hypertension, impaired fasting glucose, elevated liver enzymes, etc.)
   OR -
- MILD physical symptoms patient currently not requiring medical treatment for comorbidities (dyspnea on moderate exertion, occasional aches/pains, fatigue, etc.) - OR -
- MILD obesity-related psychological symptoms and/or mild impairment of well-being (quality of life not impacted)

#### Case Example:

38 year old female with a BMI of 59.2 kg/m², borderline hypertension, mild lower back pain, and knee pain. Patient does not require any medical intervention.

Class III, Stage 1 Obesity

Table 2.2. Proposed classification of weight by BMI in adult Asians

Classification	BMI (kg/m²)	Risk of co-morbidities
Underweight	< 18.5	Low (but increased risk of other clinical problems)
Normal range	18.5-22.9	Average
Overweight:	≥ 23	
At risk	23-24.9	Increased
Obese I	25-29.9	Moderate
Obese II	≥ 30	Severe



Patient does not meet clinical criteria

for admission at this time.

Please refer to primary care

for further preventative treatment options.

#### STAGE 2

- Patient has ESTABLISHED obesity-related comorbidities requiring medical intervention
   (HTN, Type 2 Diabetes, sleep apnea, PCOS, osteoarthritis, reflux disease) - OR -
- MODERATE obesity-related psychological symptoms (depression, eating disorders, anxiety disorder) - OR -
- MODERATE functional limitations in daily activities (quality of life is beginning to be impacted)

#### Case Example:

32 year old male with a BMI of 36 kg/m² who has primary hypertension and obstructive sleep apnea.

Class II, Stage 2 Obesity

#### STAGE 3

- Patient has significant obesity-related end-organ damage (myocardial infarction, heart failure, diabetic complications, incapacitating osteoarthritis) - OR -
- SIGNIFICANT obesity-related psychological symptoms (major depression, suicide ideation) - OR -
- SIGNIFICANT functional limitations
   (eg: unable to work or complete routine activities, reduced mobility)
- SIGNIFICANT impairment of well-being (quality of life is significantly impacted)

#### Case Example:

49 year old female with a BMI of 67 kg/m² diagnosed with sleep apnea, CV disease, GERD, and suffered from stroke. Patient's mobility is significantly limited due to osteoarthritis and gout.

Class III, Stage 3 Obesity

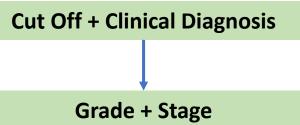
#### STAGE 4

- SEVERE (potential end stage) from obesity-related comorbidities - OR -
- SEVERELY disabling psychological symptoms OR -
- SEVERE functional limitations

#### Case Example:

45 year old female with a BMI of 54 kg/m² who is in a wheel chair because of disabling arthritis, severe hyperpnea, and anxiety disorder.

Class III, Stage 4 Obesity





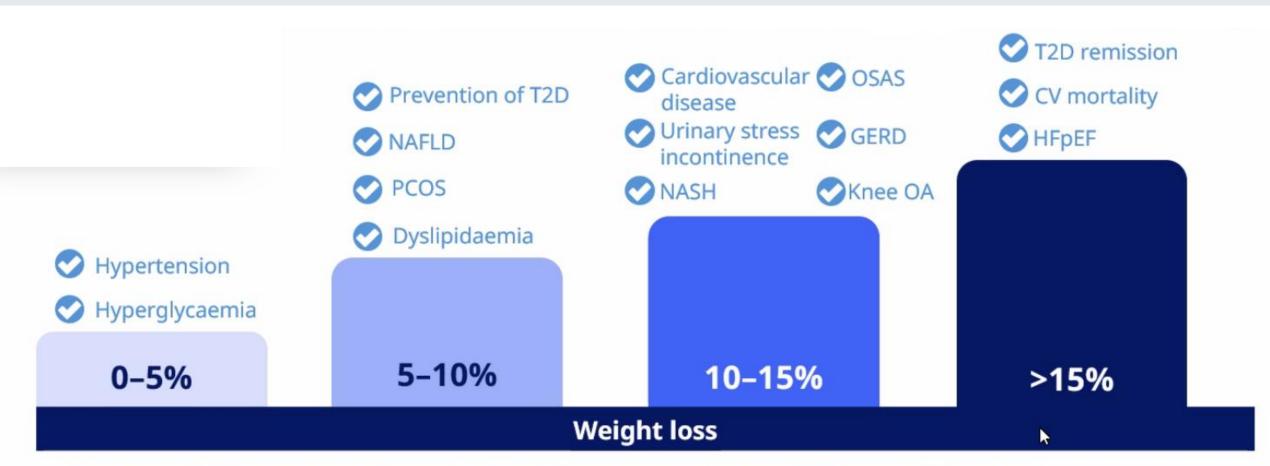


Sharma AM & Kushner RF, Int J Obes 2009

# Weight Loss and Health Improvement







CV, cardiovascular; GERD, gastro-oesophageal reflux disease; HFpEF, heart failure with preserved ejection fraction; NAFLD, non-alcoholic fatty liver disease; NASH, non-alcoholic steatohepatitis; OA: osteoarthritis; OSAS, obstructive sleep apnoea syndrome; PCOS, polycystic ovary syndrome; TG, triglycerides.

Garvey WT et al. Endocr Pract 2016;22(Suppl. 3):1–203; Look AHEAD Research Group. Lancet Diabetes Endocrinol 2016;4:913–21; Lean ME et al. Lancet 2018;391:541–51; Benraoune F and Litwin SE. Curr Opin Cardiol 2011;26:555–61; Sundström J et al. Circulation 2017;135:1577–85.

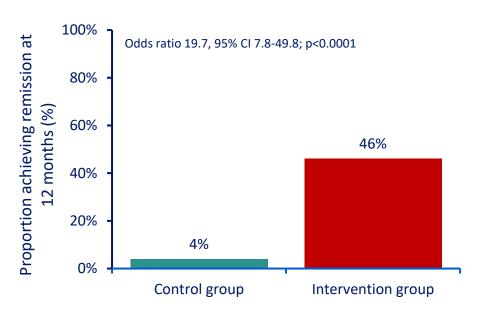
# Weight loss and T2D remission: DiRECT Trial

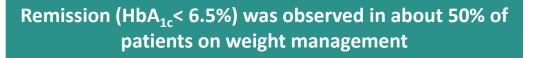
**Open-label cluster-randomised trial** 





# Weight management programme (intervention) vs best-practice care (control) in patients with T2D (no insulin; BMI of 27–45 kg/m²)



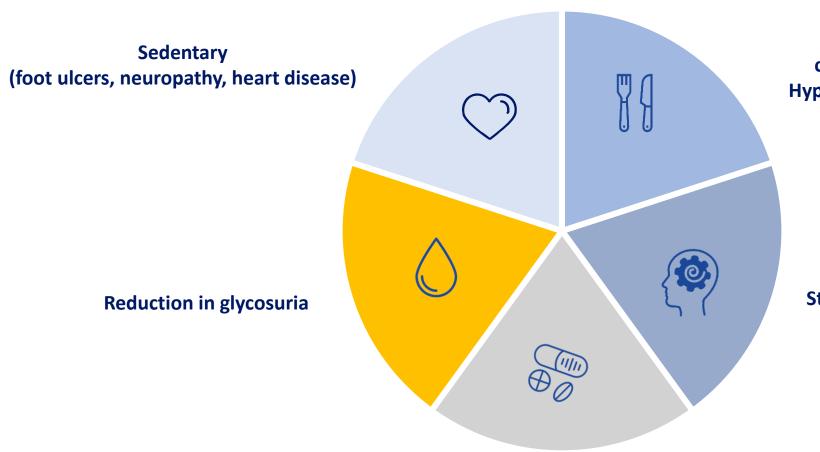




Remission rates increased with greater weight loss

# In people with diabetes, weight loss can be more difficult to achieve





Fear of hypoglycemia and compensatory over-eating; Hypoglycemia increases appetite

Stress burden of diabetes

Medications (insulin, sulfonylureas, TZDs)

# Weight Management Issues in Diabetes Mellitus

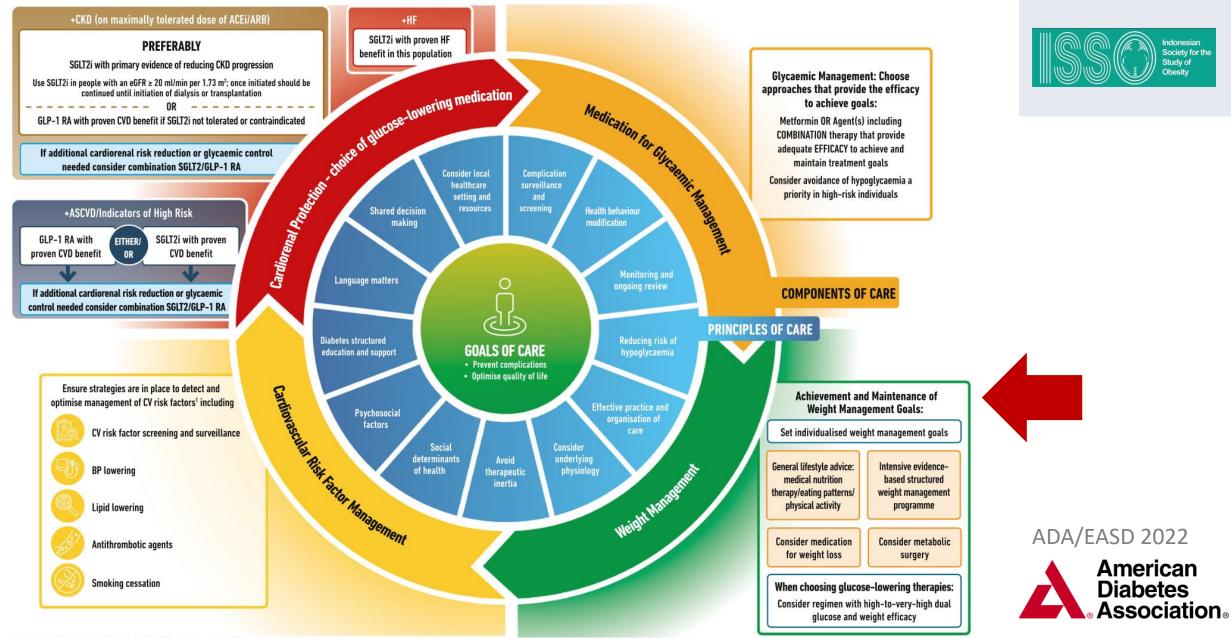




Among clinicians, weight management has been seen as the least priority in comparison to glycaemic control.

Most patients are also more willing to discuss their glycaemic target but not their body weight.

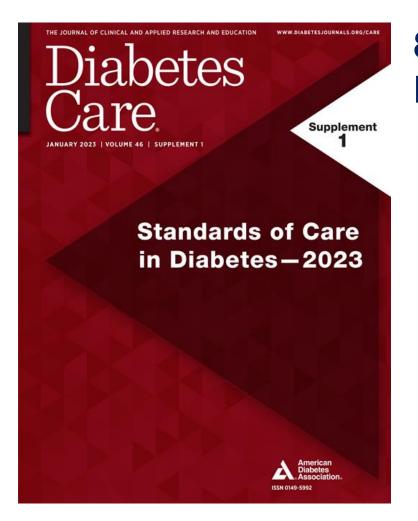
### HOLISTIC PERSON-CENTRED APPROACH TO T2DM MANAGEMENT



<sup>1 =</sup> American Diabetes Association Professional Practice Committee. 10. Cardiovascular Disease and Risk Management: Standards of Medical Care in Diabetes-2022. Diabetes Care. 2022 Jan 1;45(Suppl 1):S144-74.

### **Standard of Care in Diabetes**





# 8. Obesity and Weight Management for the Prevention and Treatment of Type 2 Diabetes

Measure height and weight and calculate BMI & assess weight trajectory to inform treatment considerations.

Individuals with diabetes and overweight or obesity may benefit from modest or larger magnitudes of weight loss.

# **Treatment Options for Weight Management in T2DM**

Table 8.1—Treatment options for overweight and obesity in type 2 diabetes







	BMI category (kg/m²)		
Treatment	25.0–26.9 (or 23.0–24.9*)	27.0-29.9 (or 25.0-27.4*)	≥30.0 (or ≥27.5*)
Nutrition, physical activity, and behavioral counseling	†	†	†

Pharmacotherapy † †

Metabolic surgery †

<sup>\*</sup>Recommended cut points for Asian American individuals (expert opinion). †Treatment may be indicated for select motivated individuals.

# **Medications Selection**





# Consider medication's effect on weight

- When choosing glucose-lowering agents
- When choosing medications for comorbidities

# Achievement and Maintenance of Weight Management Goals:

Set individualized weight management goals

General lifestyle advice: medical nutrition therapy/eating patterns/ physical activity Intensive evidencebased structured weight management program

Consider medication for weight loss

Consider metabolic surgery

#### When choosing glucose-lowering therapies:

Consider regimen with high-to-very-high dual glucose and weight efficacy

Efficacy for weight loss

Very High:

Semaglutide, Tirzepatide

High:

Dulaglutide, Liraglutide

Intermediate:

GLP-1RA (not listed above), SGLT2i

Neutral:

DPP-4i, Metformin



# **Obesity Pharmacotherapy**





# Consider obesity pharmacotherapy

- Consider potential benefits and risks
- Assess responses
  - Effective (>5% weight loss after 3 months use) --- continue
  - Insufficient early response or safety or tolerability issues consider discontinuation and evaluate alternative

# Weight Loss by Anti Obesity Medications





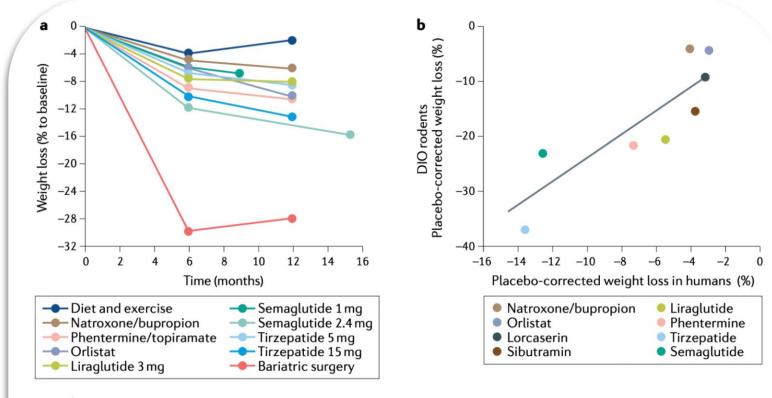


Fig. 3 | **Body weight loss by AOMs in humans and rodents.** Body weight loss achieved through lifestyle changes, currently approved anti-obesity medications (AOMs) and bariatric surgery (part **a**) and correlation of drug-induced body weight loss in rodents and humans (part **b**). Data in panel **a** refer to liraglutide 3 mg (REF. <sup>176</sup>), orlistat <sup>289</sup>, naltrexone/bupropion <sup>292</sup>, phentermine/topiramate <sup>291</sup>, semaglutide 1 mg (REF. <sup>125</sup>), semaglutide 2.4 mg (REF. <sup>38</sup>) and tirzepatide (5 and 15 mg) <sup>126</sup>. Data in panel **b** refer to naltrexone/bupropion <sup>39,295</sup>, orlistat <sup>39,296</sup>, lorcaserin <sup>39,297</sup>, sibutramine <sup>154,298</sup>, liraglutide <sup>39,299</sup>, ohentermine <sup>121,145</sup>, semaglutide <sup>38,123</sup> and tirzepatide <sup>122,127</sup>.

Nature. March 2022

# **Take Home Messages**







 Obesity is one of the important determinants of diabetes mellitus and cardiovascular diseases



 Weight reduction in diabetes may halt diabetes progression, or even promote diabetes remission. It may also lead to a reduction in CVDs.



 Weight Management in diabetes is important. A collaborative approach is needed to achieve optimal weight loss

# **THANK YOU**







