

# OVERVÆGTS-BEHANDLINGS MANGE ANSIGTER

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Institut for Idræt og Ernæring, Københavns Universitet

Nationalt Center for overvægt

# INTERESSEKONFLIKTER

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▶ Foredrag/undervisning (sidste 3 år):

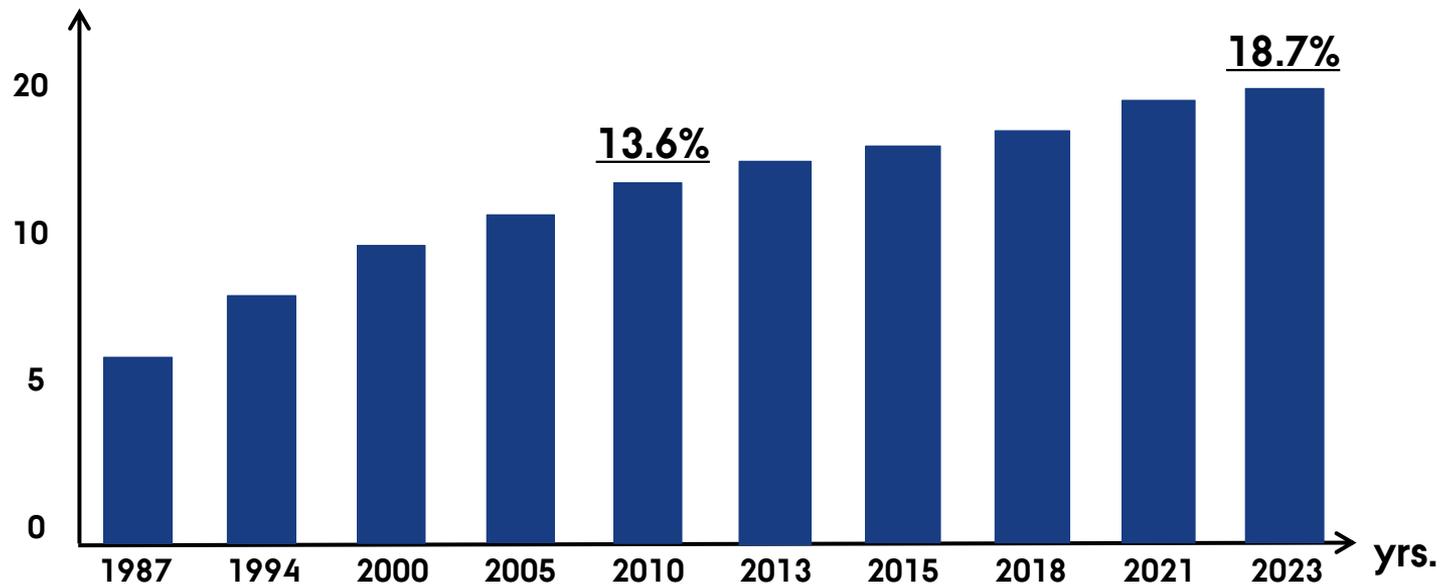
Böhringer Ingelheim, Eli Lilly, Landbrug & Fødevarer, Merck, Nestlé & Novo Nordisk

▶ Forskningsbevillinger (sidste 3 år):

Aarhus Universitet, Danske Regioner, Novo Nordisk Fonden & Sygeforsikringen danmark

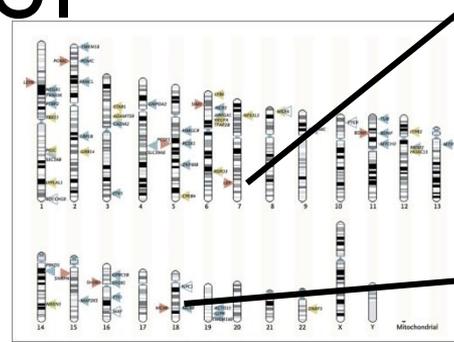
# SVÆR OVERVÆGT DANMARK (BMI>30)

Obesity (%)



# UDVIKLING AF SVÆR OVERVÆGT

- Genes
- Factors during pregnancy
- Structural/socioeconomic factors
- Psychological factors
- High caloric food intake
- Inactivity/reduced activity
- .....?



Wang Z et al, Nat Gen 2022



*Lep*

Farooqi S et al. JCI 2002

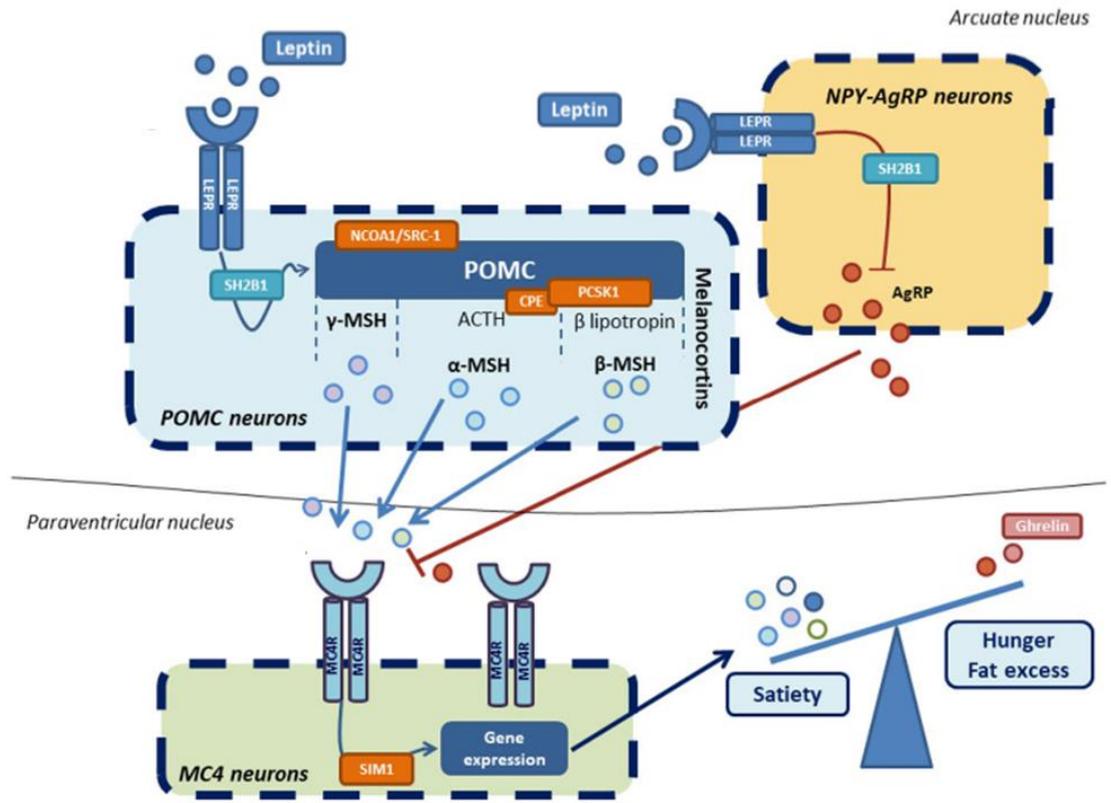
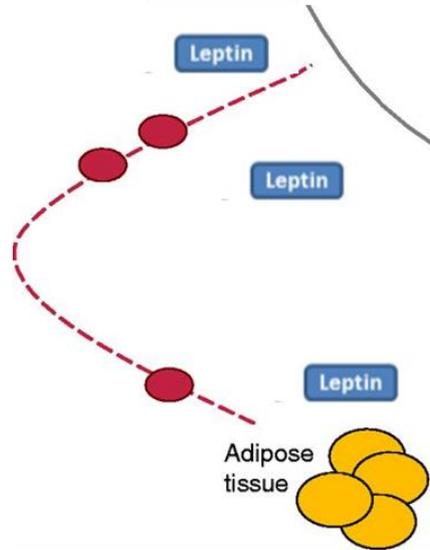


*MC4R*

Bruun JM et al  
Ugeskr Laeger 2004



# FEDT OG HJERNE



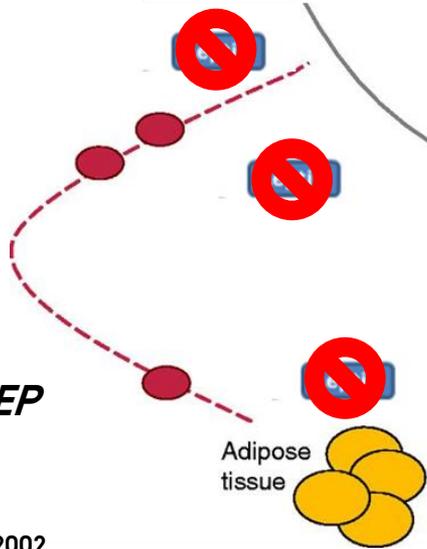
Mainieri F, Eur J Ped 2023

# FEDT OG HJERNE

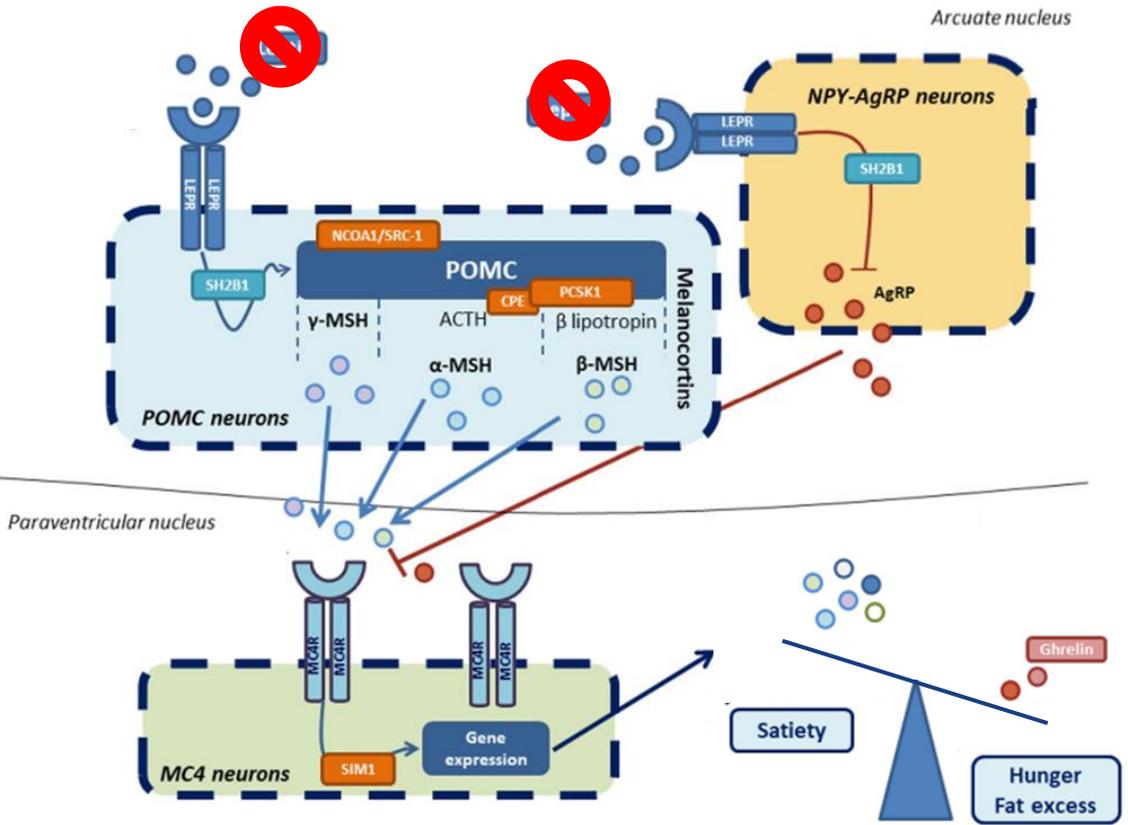


**LEP**

Farooqi S et al. JCI 2002



Adipose tissue



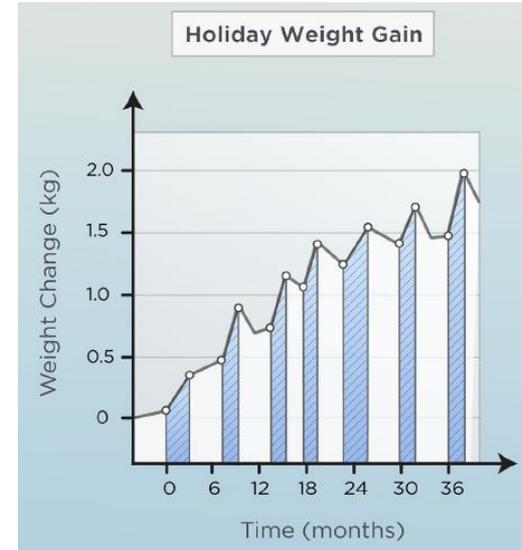
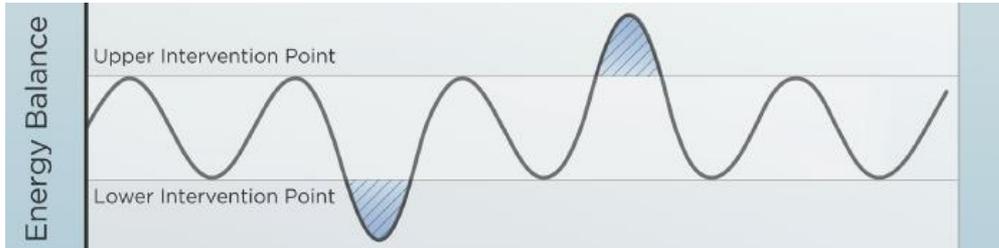
Mainieri F, Eur J Ped 2023

# VÆGTUDVIKLING I ET ALMINDELIGT LIV....

UNSOLVED MYSTERY

## The unidentified hormonal defense against weight gain

Jens Lund , Camilla Lund , Thomas Morville , Christoffer Clemmensen \*



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**”The genetic background loads the gun,  
but the environment pulls the trigger”**

Swinburn *et al.* (2011). The global obesity pandemic: shaped by global drivers and local environment

# FØLGESYGDOMME

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- ▶ Især central (abdominal) svær overvægt:

Type 2 diabetes

Hypertension/CVD

Søvnapnø

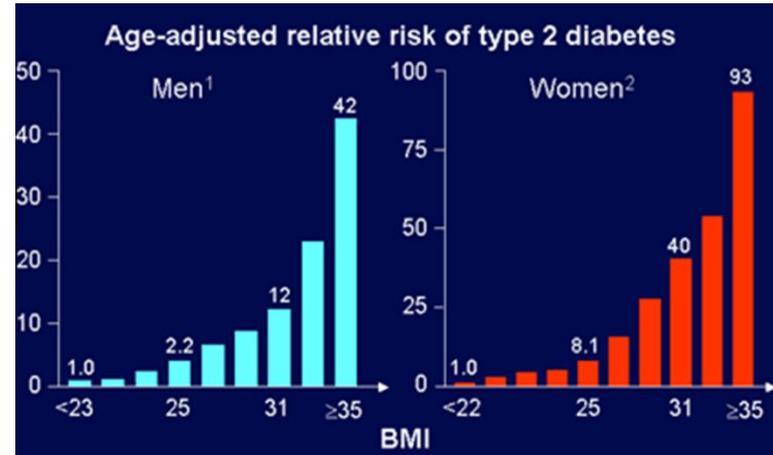
Fedtlever (MAFLD/MASH)

PCOS

Cancer

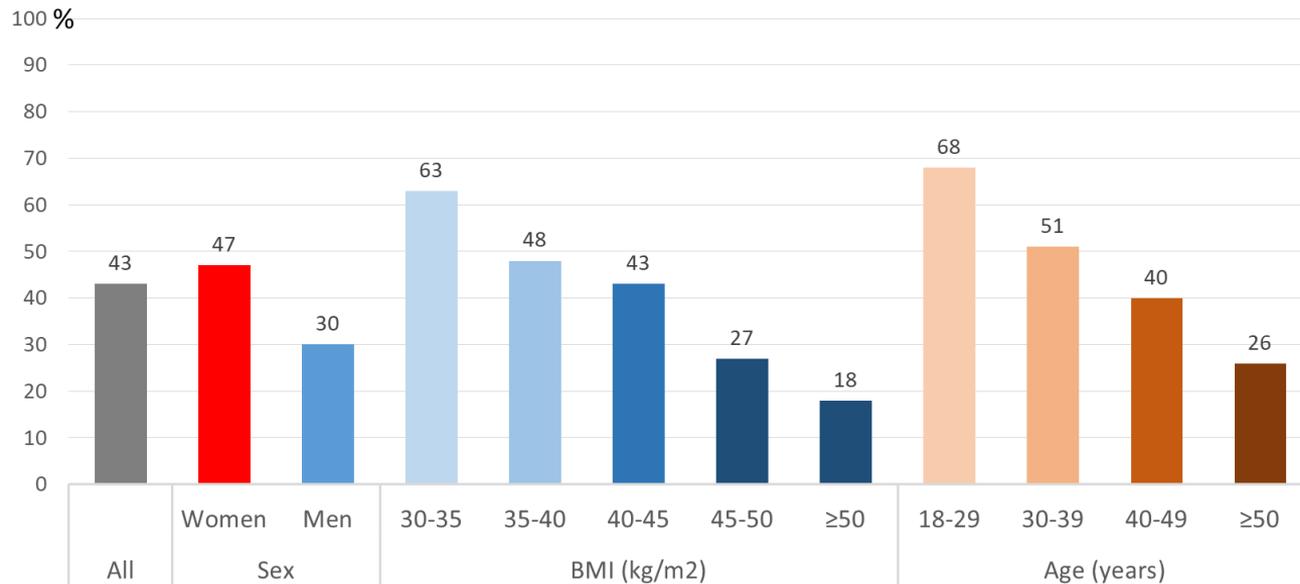
Muskel-skelet lidelser

....for tidlig død



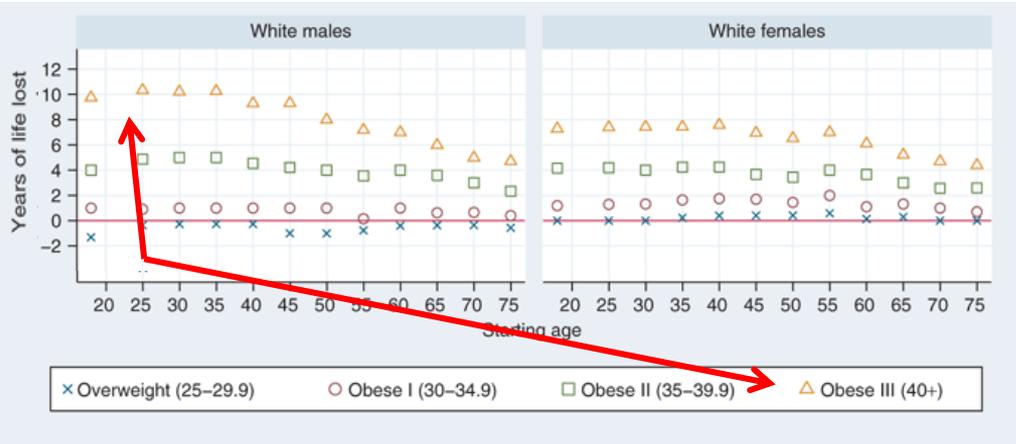
Chan JM, Diabetes Care 1994<sup>1</sup> and Colditz G, Ann Intern Med 1995<sup>2</sup>

# ...MEN HØJT BMI ER *IKKE* LIG MED SYGDOM

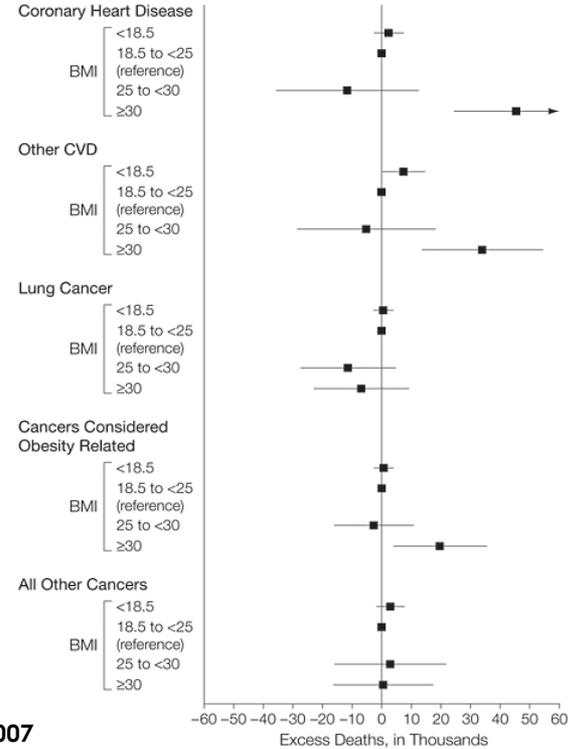


\*Ikke T2D, prædiabetes, øget leverstivhed eller obstruktiv søvnapnø

# MORTALITET



Finkelstein EA, Obesity 2010



Flegal KM, JAMA 2007

# KAN VI FOREBYGGE?



Received: 26 October 2022 | Revised: 18 August 2023 | Accepted: 9 September 2023

DOI: 10.1111/obr.13641

## REVIEW

Paediatric Body weight - Policy

OBESITY  
Reviews WILEY

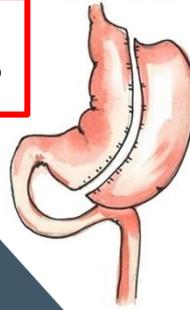
**A literature review of evidence for primary prevention of overweight and obesity in healthy weight children and adolescents: A report produced by a working group of the Danish Council on Health and Disease Prevention**

Nanna J. Olsen<sup>1</sup>  | Jane N. Østergaard<sup>2</sup> | Lise G. Bjerregaard<sup>3</sup> | Teresa V. Høy<sup>4</sup> |  
Lene Kierkegaard<sup>4</sup> | Kim F. Michaelsen<sup>5</sup> | Thorkild I. A. Sørensen<sup>6</sup> |  
Morten K. Grønæk<sup>4</sup> | Jens M. Bruun<sup>2,7,8</sup>  | Berit L. Heitmann<sup>1,9,10</sup>

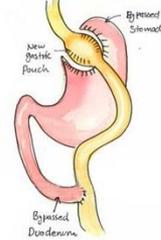
# BEHANDLING

Weight loss ~30%

Gastric Sleeve



Gastric Bypass



Weight loss ~5%

Weight loss 10-20%

Kirurgi

Medicin

Livsstil

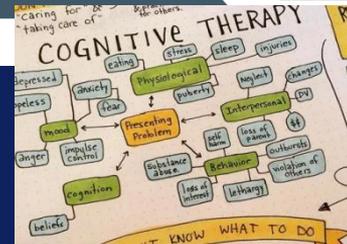
(Diæt, fysisk aktivitet, adfærdspåvirkning)

**Mysimba<sup>®</sup> 8 mg/90 mg**  
prolonged-release tablets

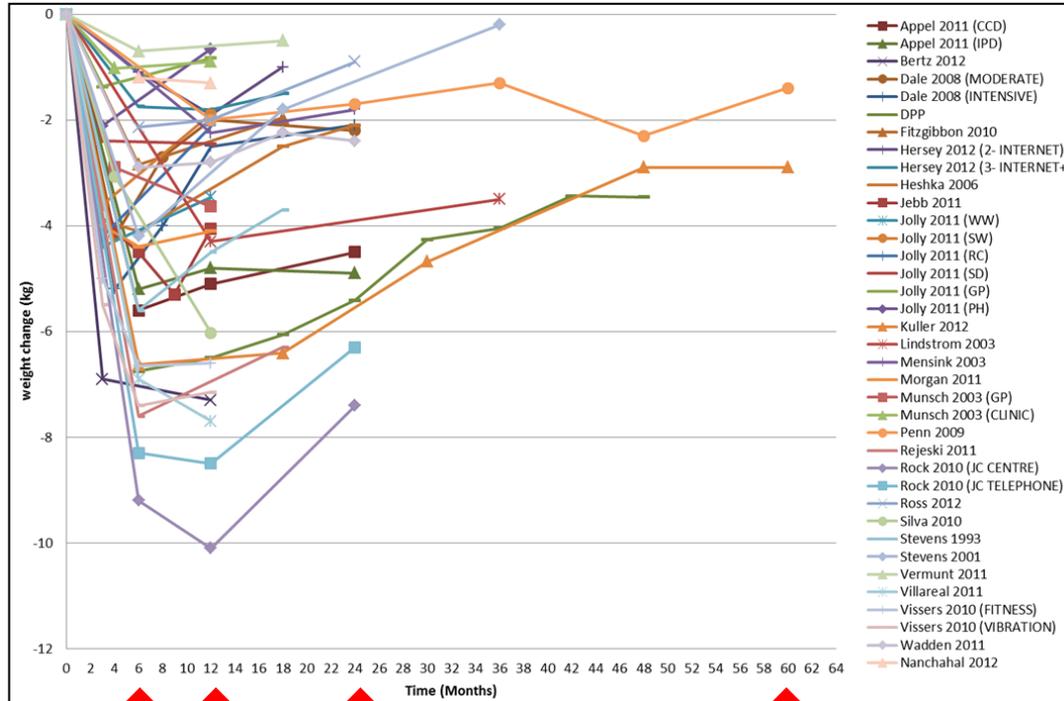
naltrexone hydrochloride / bupropion hydrochloride  
112 prolonged-release tablets



once weekly  
**mounjaro<sup>®</sup>**  
(tirzepatide) injection 0.5 mL  
2.5 mg | 5 mg | 7.5 mg | 10 mg | 12.5 mg | 15 mg

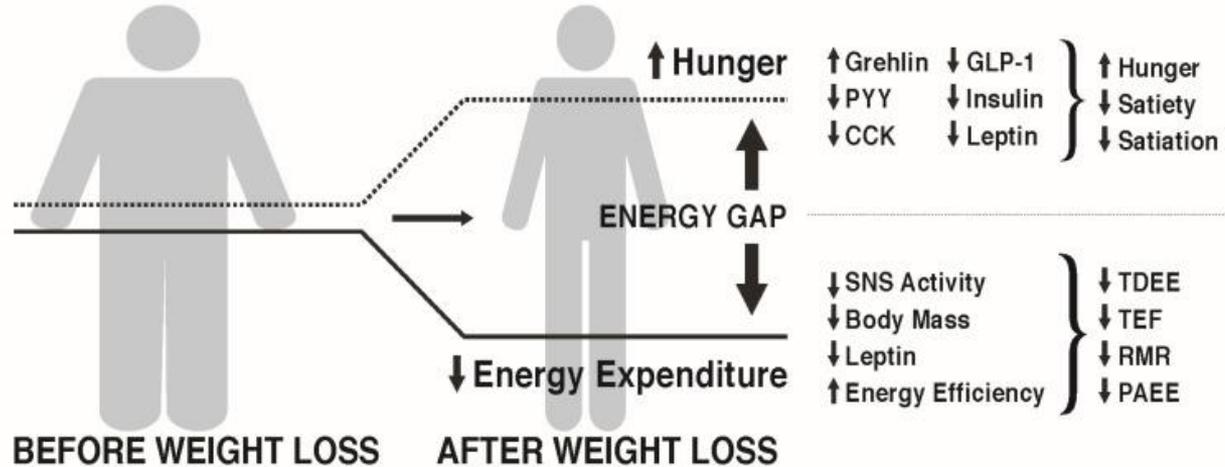


# LIVSSTILS INTERVENTIONER



By courtesy of professor Jebb

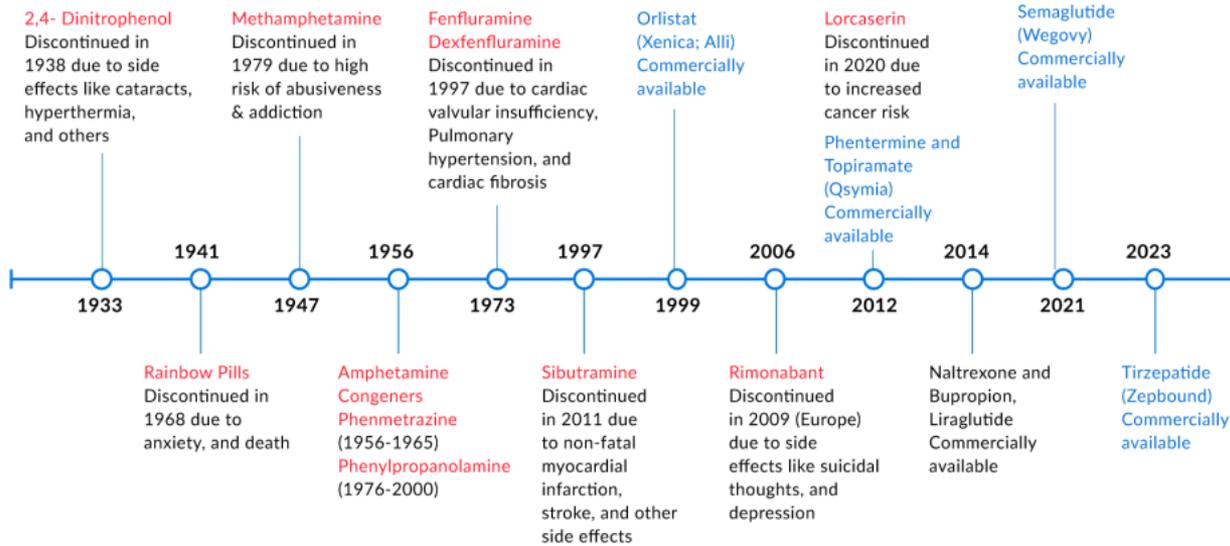
# VÆGTTAB OG VÆGTVEDLIGEHOEDSE



Melby CL, Nutrients 2017

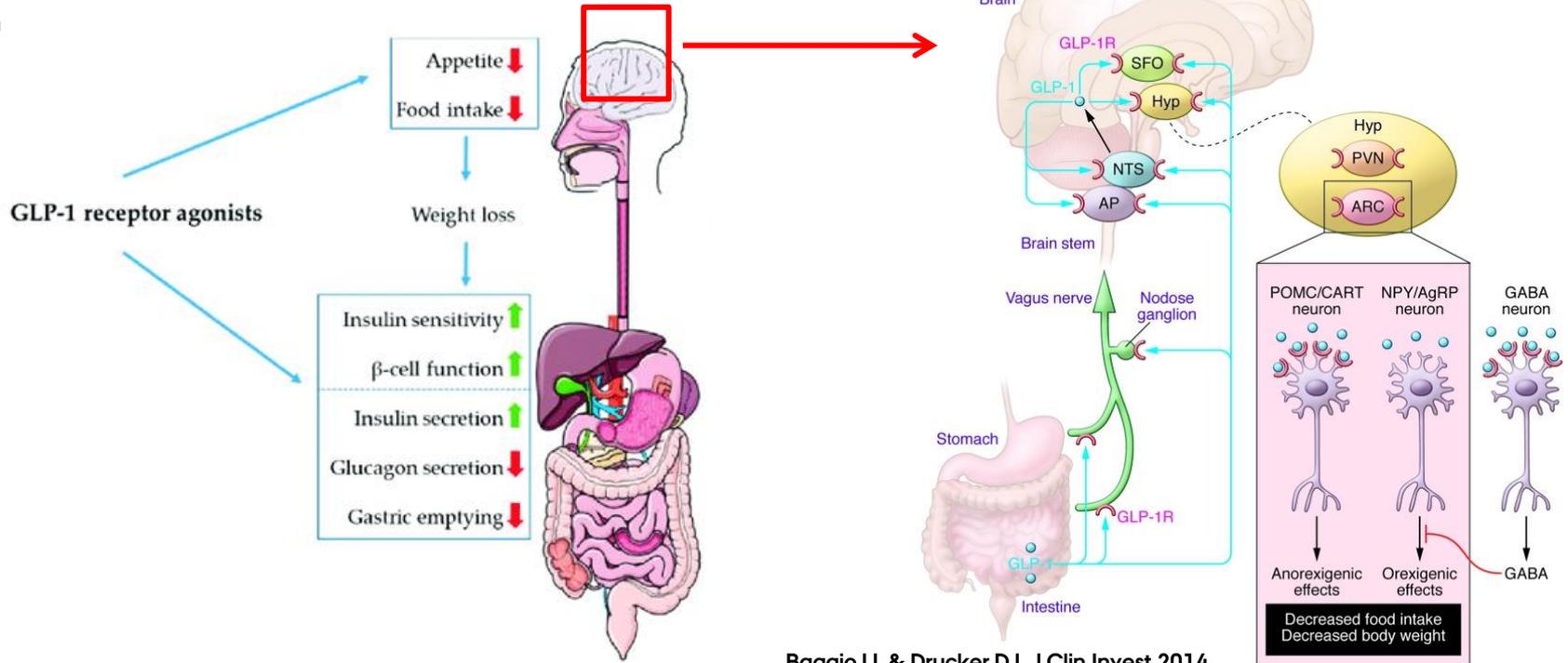
# FARMAKOLOGI

## Obesity Drugs Timeline



● Red color indicates discontinued drugs ● Blue color indicates commercially available drugs

# GLP1-RA



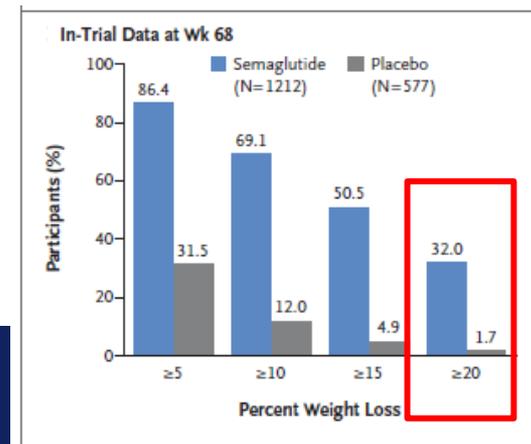
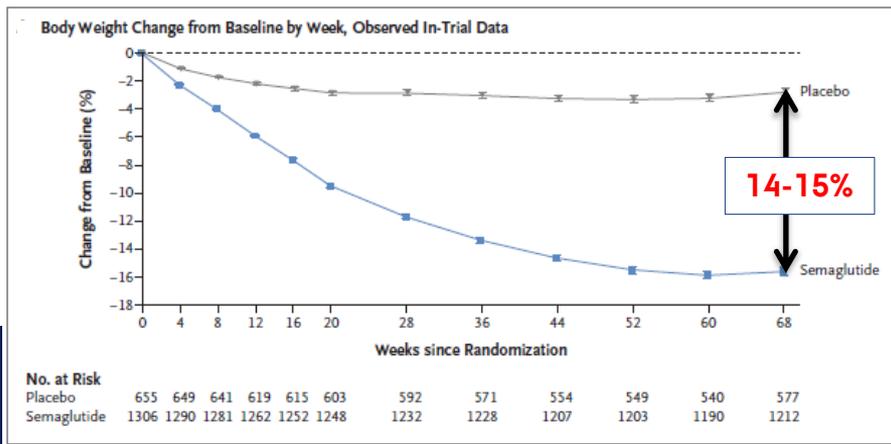
Baggio LL & Drucker DJ, J Clin Invest 2014

# WEGOVY (SEMAGLUTID)

ORIGINAL ARTICLE

## Once-Weekly Semaglutide in Adults with Overweight or Obesity

John P.H. Wilding, D.M., Rachel L. Batterham, M.B., B.S., Ph.D., Salvatore Calanna, Ph.D., Melanie Davies, M.D., Luc F. Van Gaal, M.D., Ph.D., Ildiko Lingvay, M.D., M.P.H., M.S.C.S., Barbara M. McGowan, M.D., Ph.D., Julio Rosenstock, M.D., Marie T.D. Tran, M.D., Ph.D., Thomas A. Wadden, Ph.D., Sean Wharton, M.D., Pharm.D., Koutaro Yokote, M.D., Ph.D., Niels Zeuthen, M.Sc., and Robert F. Kushner, M.D., for the STEP 1 Study Group\*



Wilding JPH, NEJM 2021

# OPHØR MED WEGOVY.....?

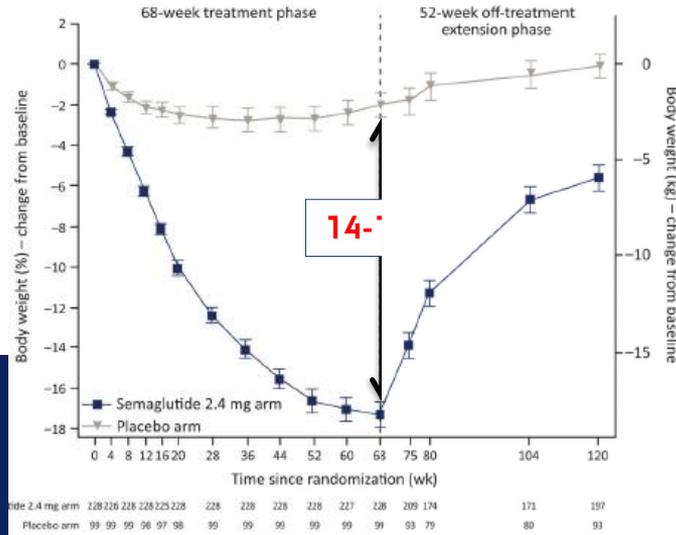
Received: 1 February 2022 | Revised: 7 April 2022 | Accepted: 18 April 2022  
DOI: 10.1111/dom.14725

ORIGINAL ARTICLE

WILEY

## Weight regain and cardiometabolic effects after withdrawal of semaglutide: The STEP 1 trial extension

John P. H. Wilding D.M.<sup>1</sup> | Rachel L. Batterham MBBS<sup>2,3,4</sup> |  
Melanie Davies M.D.<sup>5,6</sup> | Luc F. Van Gaal M.D.<sup>7</sup> | Kristian Kandler M.D.<sup>8</sup> |  
Katerina Konakli PhD<sup>8</sup> | Ildiko Lingvay M.D.<sup>9</sup> | Barbara M. McGowan M.D.<sup>10</sup> |  
Tugce Kalayci Oral MD<sup>8</sup> | Julio Rosenstock M.D.<sup>11</sup> |  
Thomas A. Wadden Ph.D.<sup>12</sup> | Sean Wharton M.D.<sup>13</sup> | Koutaro Yokote M.D.<sup>14</sup> |  
Robert F. Kushner M.D.<sup>15</sup> | STEP 1 Study Group



Wilding JPH, DOM 2022

# MOUNJARO (TIRZEPATID)

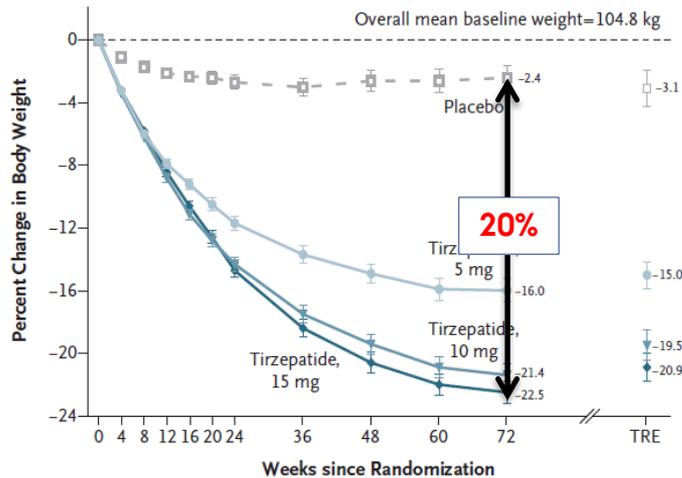
The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

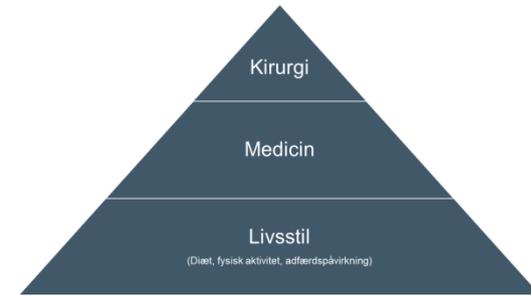
## Tirzepatide Once Weekly for the Treatment of Obesity

Ania M. Jastreboff, M.D., Ph.D., Louis J. Aronne, M.D.,  
 Nadia N. Ahmad, M.D., M.P.H., Sean Wharton, M.D., Pharm.D.,  
 Lisa Connery, M.D., Breno Alves, M.D., Arihiro Kiyosue, M.D., Ph.D.,  
 Shuyu Zhang, M.S., Bing Liu, Ph.D., Mathijs C. Bunck, M.D., Ph.D.,  
 and Adam Stefanski, M.D., Ph.D., for the SURMOUNT-1 Investigators\*

Percent Change in Body Weight by Week (efficacy estimand)



Participants Who Met Weight-Reduction Targets (efficacy estimand)



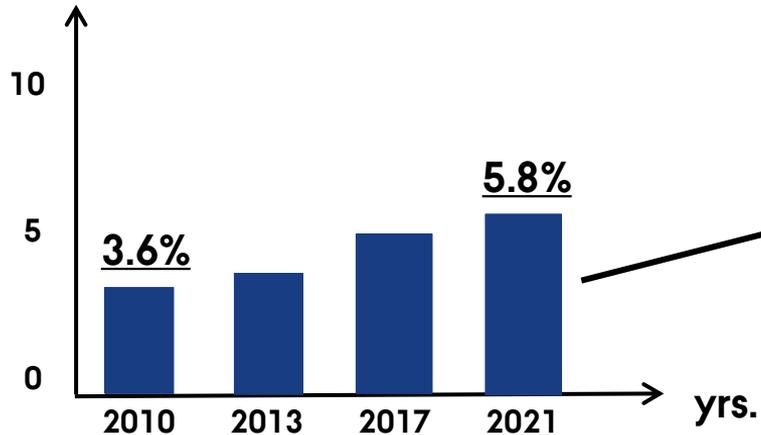
Jastreboff AM, NEJM 2022



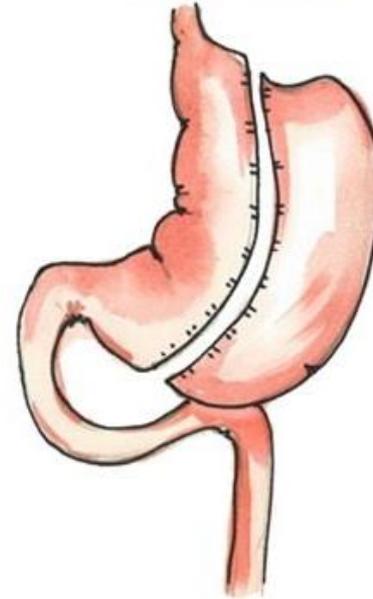
# SVÆR OVERVÆGT KLASSE II+III (BMI>35)

~1400-1600/år i Danmark

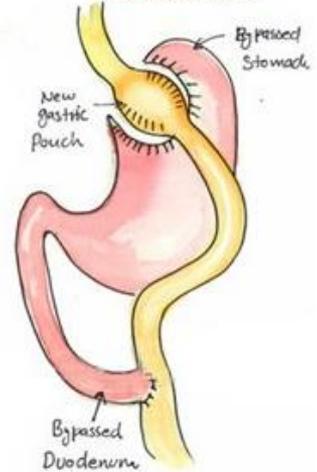
Obesity (%)



Gastric Sleeve

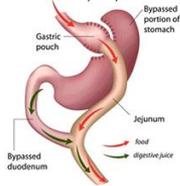


Gastric Bypass

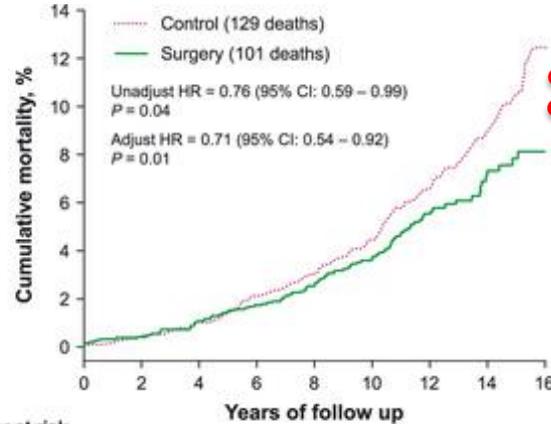
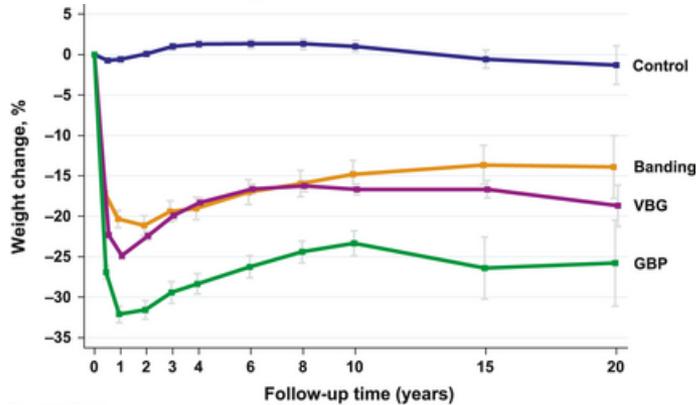
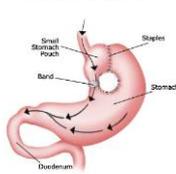


# SOS

Roux-en-Y Gastric Bypass (RNY)



Vertical Banded Gastroplasty



Number at risk

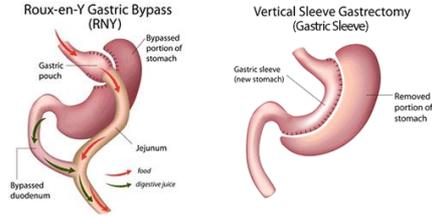
	2010	2001	1987	1821	1590	1260	760	422	169
Surgery	2037	2027	2016	1842	1455	1174	749	422	156
Control	2037	2027	2016	1842	1455	1174	749	422	156

Table 2. Cause of Death.\*

Variable	Surgery Group (N = 2010)	Control Group (N = 2037)
	no. of subjects	
<b>Cardiovascular condition</b>		
Any event	43	52
Cardiac	35	44
Myocardial infarction	13	25
Heart failure	2	5
Sudden death	20	14
Stroke	6	6
Intracerebral hemorrhage	2	4
Infarction	1	2
Subarachnoid bleeding	3	0
Other	2	3
Aortic aneurysm	1	2
Aortic thrombosis	0	1
Diabetic gangrene	1	0
<b>Noncardiovascular condition</b>		
Any event	58	76
Cancer	29	47
Meningioma	0	1
Infection	12	3
Thromboembolic disease	5	7
Pulmonary embolism	4	7
Vena caval thrombosis	1	0
Other	12	18
Total no. of deaths	101	129

Sjöström L, J Intern Med 2012 & N Eng J Med 2007

# T2D (3 ÅR EFTER)

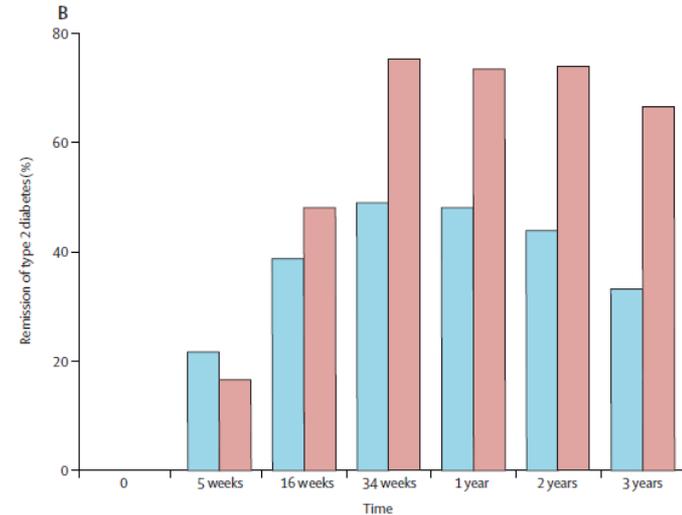
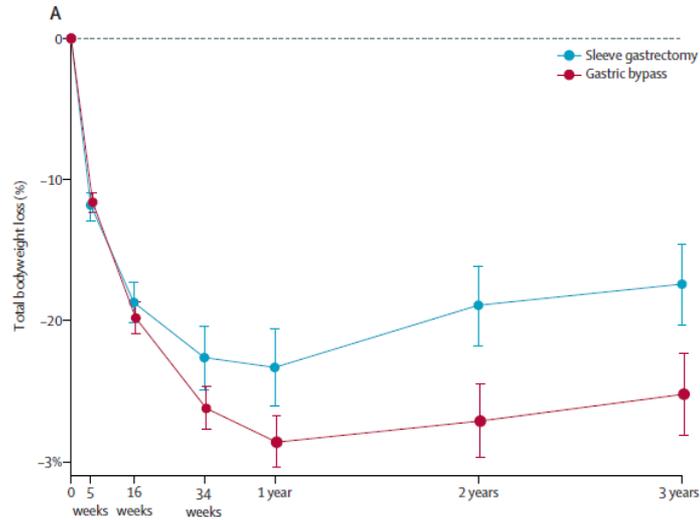


## Patient-reported outcomes, weight loss, and remission of type 2 diabetes 3 years after gastric bypass and sleeve gastrectomy (Oseberg); a single-centre, randomised controlled trial



Marius Svanevik\*, Jolanta Lorentzen\*, Heidi Borgeraas, Rune Sandbu, Birgitte Seip, Asle W Medhus, Jens K Hertel, Ronette L Kolotkin, Milada C Småstuen, Dag Hofsoe†, Joran Hjelmæsæth†

Lancet Diabetes Endocrinol  
2023; 11: 555-66



# LIDT AT GÅ HJEM PÅ

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- ▶ Forebyggelse er svær (kan vi overhovedet forebygge?)
- ▶ Livsstilintervention virker, så længe livsstilen er lagt om....
- ▶ Farmakologisk behandling kan reducere kropsvægten med 20%, men virker kun så længe man tager medicinen
- ▶ Bariatrisk kirurgi forbedrer metabolisme (T2D) og reducerer mortalitet
- ▶ Mere forskning....