

Obesity beyond BMI - Obesities, Clinical Obesity

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Disclosure

**The Speaker has no conflict of interest
regarding this presentation**

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- **Body Mass Index (BMI) : a standard measure of Adiposity?**
- **Issues in Obesity, originated from BMI**
- **Obesities, beyond BMI**
- **Clinical Obesity – Obesity addressed in clinical practice**

57 years old Man

- Height 1.81 m
- Weight 87 Kg
- BMI 36.5 Kg/m²
- Waist Circ 100 cm
- Family History of
 - Acute MI, T2DM
 - Stroke
 - Dementia
 - Parkinsonism
 - Colon cancer
- Non-smoker
- Blood pressure 122/78 mmHg
- Glucose 89 mg/dl
- HbA1c 5.6%
- Lipid profile 206/146/69/108 mg/dl
- Mild fatty liver on abdominal us
- Past History of
 - Hepatitis B
 - Pulmonary Tuberculosis
 - Herpes Zoster
 - Exercise induced Asthma
 - Atopic dermatitis
- 10 units of alcohol once a week

Body Mass Index (BMI) : a standard measure of Adiposity?



The definition of Obesity

- a " chronic, relapsing, multi-factorial, neurobehavioral disease, wherein an increase in body fat promotes adipose tissue dysfunction and abnormal fat mass physical forces, resulting in adverse metabolic, biomechanical, and psychosocial health consequences."

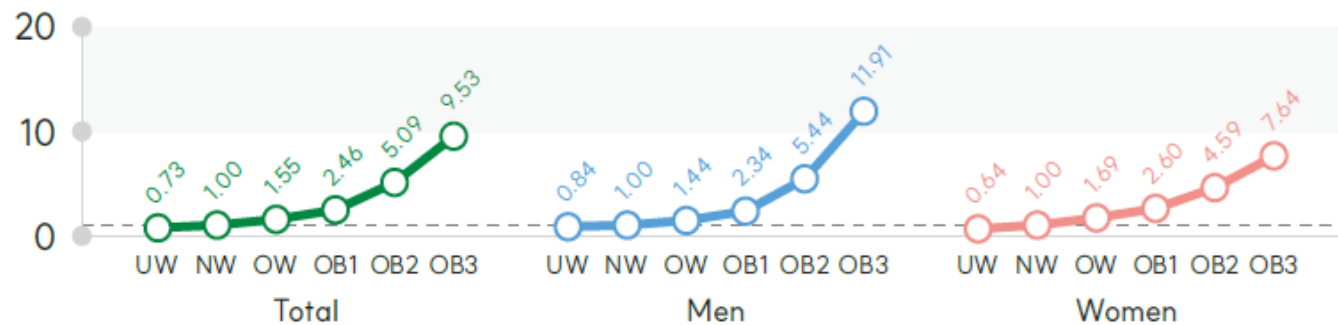
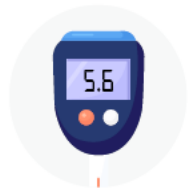
The Obesity Medicine Association,

The definition of Obesity

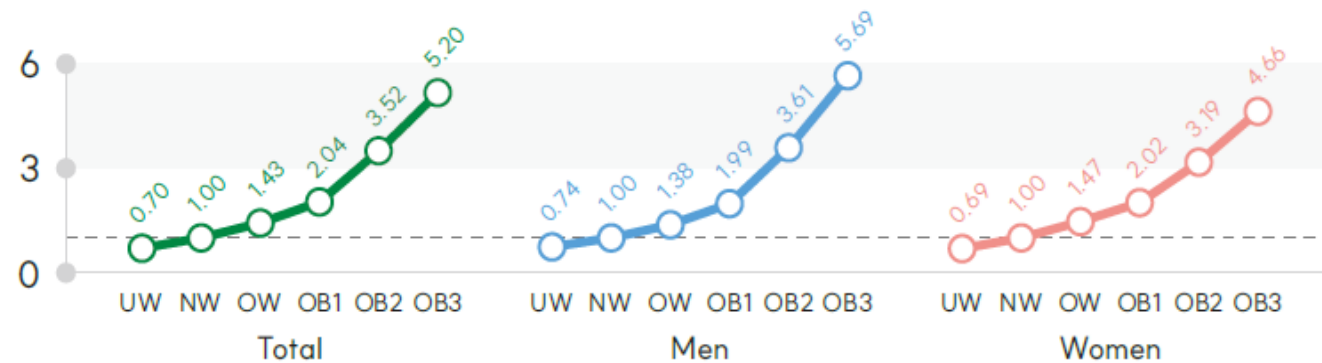
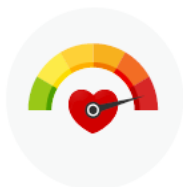
$$\frac{\text{Weight in Kilogram}}{(\text{Height in Meter})^2} \geq \mathbf{30} \text{ Or } \mathbf{25}$$

Risk of chronic disease incidence by obesity class based on BMI

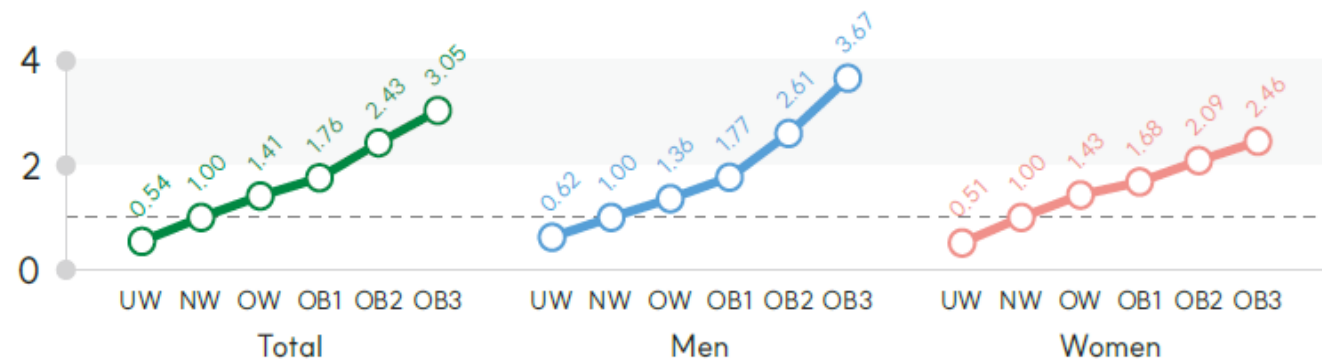
Type 2 diabetes



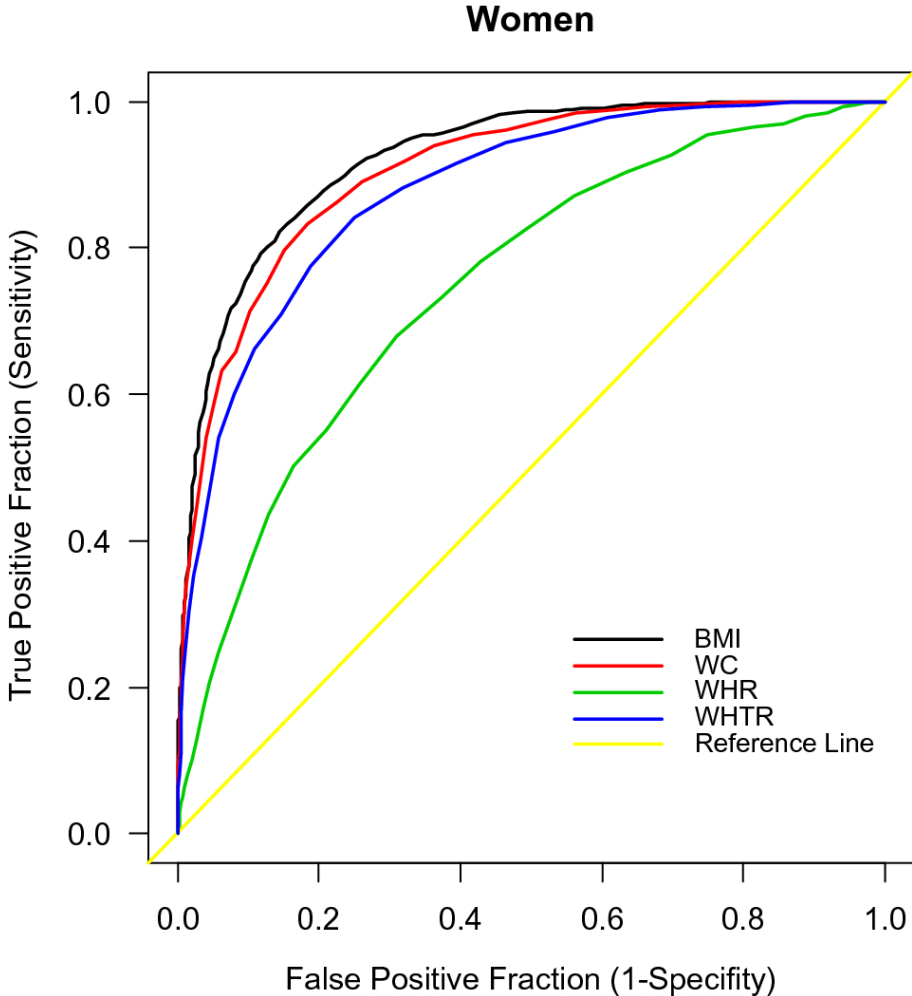
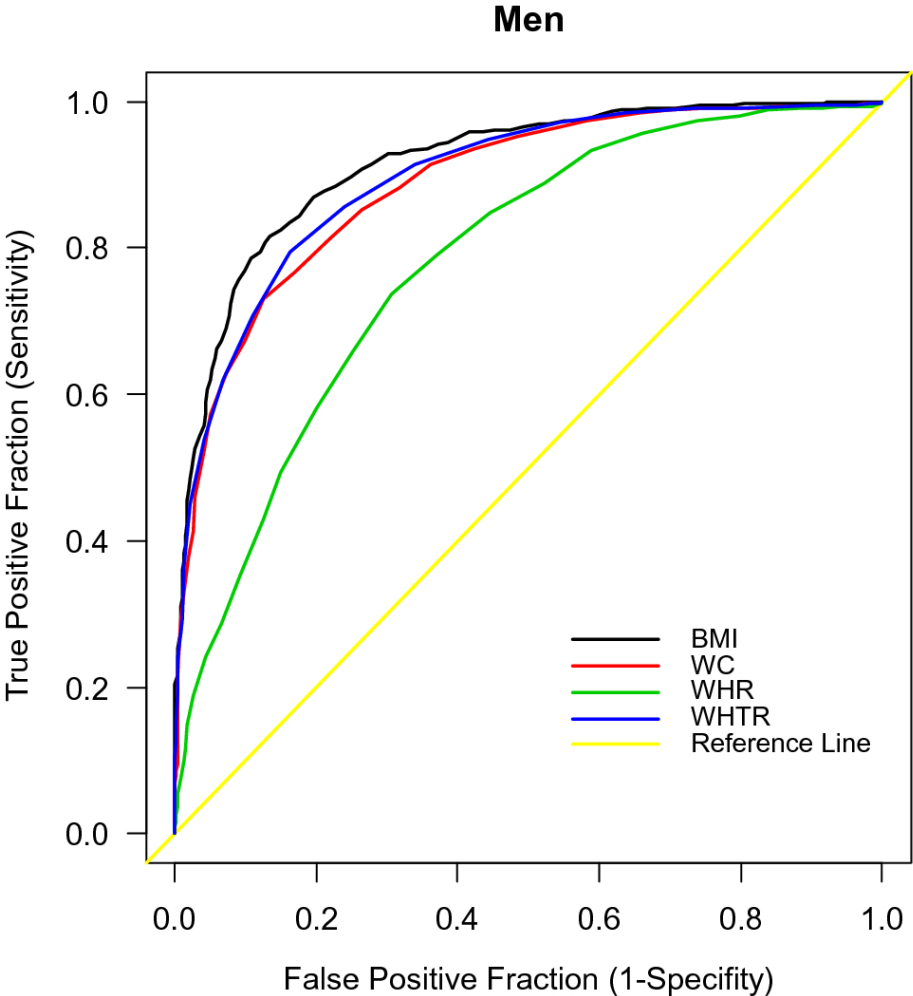
Hypertension



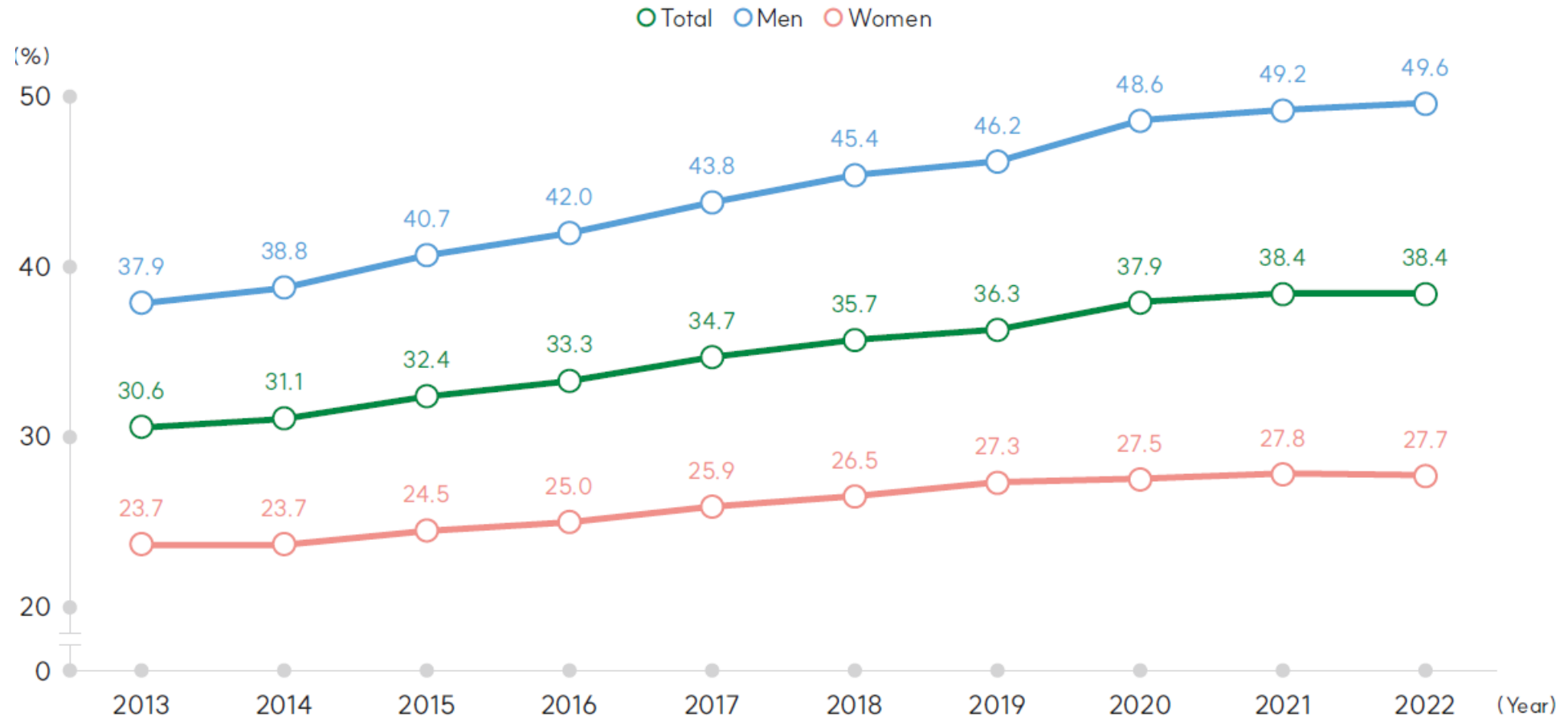
Dyslipidemia



ROC curves for BMI, WC, WHR, and WHTR in screening for cardiovascular risk factors



Prevalence of obesity in Korea during last 10 years



The diagnosis and classification of Obesity

WEIGHT		95	100	105	110	115	120	125	130	135	140	145	150	155	160	165	170	175	180	185	190	195	200	205	210	215	220	225	230	235	240	245
HEIGHT	5'0"	19	20	21	21	22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42	43	44	45	46	47	48
	5'1"	18	19	20	21	22	23	24	25	26	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42	43	43	44	45	46
	5'2"	17	18	19	20	21	22	23	24	25	26	27	27	28	29	30	31	32	33	34	35	36	37	37	38	39	40	41	42	43	44	45
	5'3"	17	18	19	19	20	21	22	23	24	25	26	27	27	28	29	30	31	32	33	34	35	35	36	37	38	39	40	41	42	43	43
	5'4"	16	17	18	19	20	21	21	22	23	24	25	26	27	27	28	29	30	31	32	33	33	34	35	36	37	38	39	39	40	41	42
	5'5"	16	17	17	18	19	20	21	22	22	23	24	25	26	27	27	28	29	30	31	32	32	33	34	35	36	37	37	38	39	40	41
	5'6"	15	16	17	18	19	19	20	21	22	23	23	24	25	26	27	27	28	29	30	31	31	32	33	34	35	36	36	37	38	39	40
	5'7"	15	16	16	17	18	19	20	20	21	22	23	23	24	25	26	27	27	28	29	30	31	31	32	33	34	34	35	36	37	38	38
	5'8"	14	15	16	17	17	18	19	20	21	21	22	23	24	24	25	26	27	27	28	29	30	30	31	32	33	33	34	35	36	36	37
	5'9"	14	15	16	16	17	18	18	19	20	21	21	22	23	24	24	25	26	27	27	28	29	30	30	31	32	32	33	34	35	35	36
	5'10"	14	14	15	16	16	17	18	19	19	20	21	22	22	23	24	24	25	26	27	27	28	29	29	30	31	32	32	33	34	34	35
	5'11"	13	14	15	15	16	17	17	18	19	20	20	21	22	22	23	24	24	25	26	26	27	28	29	29	30	31	31	32	33	33	34
	6'0"	13	14	14	15	16	16	17	18	18	19	20	20	21	22	22	23	24	24	25	26	26	27	28	28	29	30	31	31	32	33	33
	6'1"	13	13	14	15	15	16	16	17	18	18	19	20	20	21	22	22	23	24	24	25	26	26	27	28	28	29	30	30	31	32	32
	6'2"	12	13	13	14	15	15	16	17	17	18	19	19	20	21	21	22	22	23	24	24	25	26	26	27	28	28	29	30	30	31	31
	6'3"	12	12	13	14	14	15	16	16	17	17	18	19	19	20	21	21	22	22	23	24	24	25	26	26	27	27	28	29	29	30	31
	6'4"	12	12	13	13	14	15	15	16	16	17	18	18	19	19	20	21	21	22	23	23	24	24	25	26	26	27	27	28	29	29	30

Under healthy weight: BMI <18.5

Healthy weight: BMI 18.5-24.9

Overweight: BMI 25-29.9

Obese I: BMI 30-34.9

Obese II: BMI 35-39.9

Obese III: BMI ≥ 40

Adolphe Quetelet

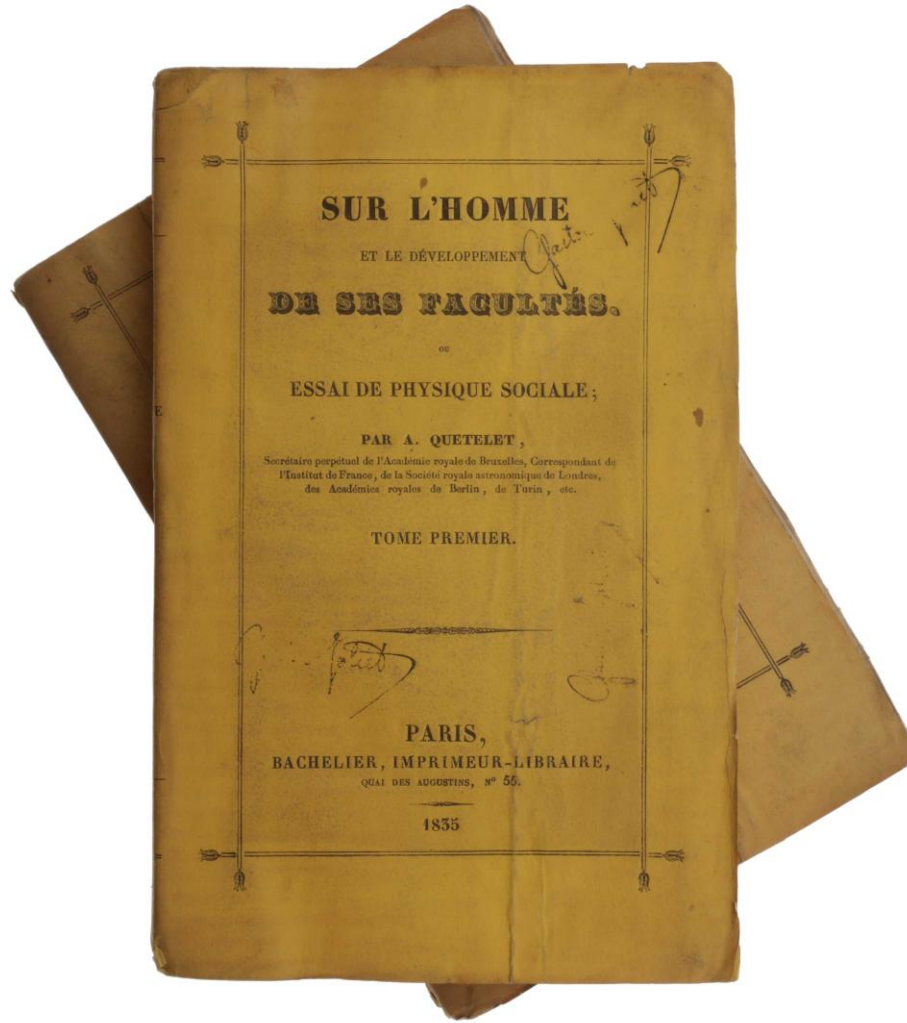


(1796 – 1874)

a Belgian
astronomer,
mathematician,
statistician
and sociologist

The founder of
anthropometry

Sur l'homme et le développement de ses facultés, ou Essai de physique sociale



A Social Treatise on Man and the Development of his Faculties

described a man's weight in relation to his height in his quest to find values for the “Average man”

1835

Standard body weight chart for Metropolitan Life insurance company in 1920s

Table of the Size and Weight of Man at Different Ages.								Table of the Size and Weight of Woman at Different Ages.							
Ages.	Size.	Weight.	Ratio of Weight to Size.	Size Observed.		Weight Observed.		Ages.	Size.	Weight.	Ratio of Weight to Size.	Size Observed.		Weight Observed.	
				Max.	Min.	Max.	Min.					Max.	Min.	Max.	Min.
	met.	kilog.		met.	met.	kilog.	kilog.		met.	kilog.		met.	met.	kilog.	kilog.
Birth,	0.496	3.20	6.19	0.532	0.438	4.50	2.34	Birth,	0.483	2.91	6.15	0.555	0.438	4.25	1.12
1 year,	0.696	10.00	14.20	0.750	0.682	11.00	9.00	1 year,	0.690	9.30	13.50	0.704	0.660	10.5	8.3
2 ..	0.797	12.00	15.00	0.824	0.730	13.50	10.50	2 ..	0.780	11.40	14.50	0.798	0.720	12.0	8.3
3 ..	0.860	13.21	15.36	0.875	0.840	13.60	12.10	3 ..	0.850	12.45	14.70	0.895	0.795	15.8	10.5
4 ..	0.932	15.07	16.32	0.965	0.840	18.20	12.50	4 ..	0.910	14.18	15.10	0.950	0.810	15.8	11.5
5 ..	0.990	16.70	16.98	1.000	0.915	18.50	14.00	5 ..	0.974	15.50	15.70	1.085	0.876	17.5	13.3
6 ..	1.046	18.04	17.44	1.115	0.960	20.40	15.80	6 ..	1.032	16.74	16.24	1.085	0.956	20.3	13.3
7 ..	1.112	20.16	18.31	1.162	1.109	24.50	17.20	7 ..	1.096	18.45	16.85	1.177	1.050	23.4	16.0
8 ..	1.170	22.26	18.92	1.260	1.120	28.50	19.00	8 ..	1.139	19.82	17.45	1.330	1.050	23.4	16.0
9 ..	1.227	24.09	19.68	1.325	1.150	29.00	22.20	9 ..	1.200	22.44	18.65	1.330	1.110	25.7	18.3
10 ..	1.282	26.12	20.37	1.325*	1.163	32.00	22.70	10 ..	1.248	24.24	19.45	1.330	1.160	28.3	20.3
11 ..	1.327	27.85	21.58	1.405	1.215	33.80	25.00	11 ..	1.275	26.25	20.60	1.385	1.160	30.8	21.6
12 ..	1.359	31.00	22.80	1.450	1.270	36.30	25.00	12 ..	1.327	30.54	23.00	1.476	1.160	42.3	21.6
13 ..	1.403	35.32	25.30	1.490	1.300	39.50	34.60	13 ..	1.336	34.65	24.50	1.580	1.160	42.8	21.6
14 ..	1.487	40.50	27.49	1.630	1.330	45.00	37.00	14 ..	1.447	38.10	25.35	1.580	1.160	51.0	32.0
15 ..	1.559	46.41	29.88	1.658	1.390	61.50	37.00	15 ..	1.475	41.30	28.10	1.638	1.160	55.2	32.0
16 ..	1.610	53.39	33.00	1.730	1.430	61.50	40.00	16 ..	1.500	44.44	29.62	1.638	1.160	57.6	32.0
17 ..	1.670	57.40	34.25	1.790	1.467	65.50	45.00	17 ..	1.544	49.08	31.75	1.688	1.224	61.6	..
18 ..	1.700	61.26	35.67	1.790	..	67.00	45.00	18 ..	1.562	53.10	34.05	1.740	..	79.9	..
19 ..	1.706	63.32	37.00	1.800	..	70.00	48.20	20 ..	1.570	54.46	34.70
20 ..	1.711	65.00	37.99	1.838	..	72.70	..	25 ..	1.577	55.08	35.26
25 ..	1.722	68.29	39.66	1.890	..	98.50	..	30 ..	1.579	55.14	35.90
30 ..	1.722	68.90	40.02	40 ..	1.555	56.65	36.50
40 ..	1.713	68.81	40.03	50 ..	1.536	58.45	38.15	..	1.444	90.5	39.8
50 ..	1.674	67.45	40.14	60 ..	1.516	56.73	37.28	..	1.436
60 ..	1.639	65.50	40.01	70 ..	1.514	53.72	35.49	..	1.431	93.8	..
70 ..	1.623	63.03	38.83	49.1	80 ..	1.506	51.52	34.21	1.701	1.408	72.5	38.0
80 ..	1.613	61.22	37.96	1.820	1.467	83.00	49.7								

The rise of Body Mass Index



Ancel Keys (1904–2004).

an American physiologist

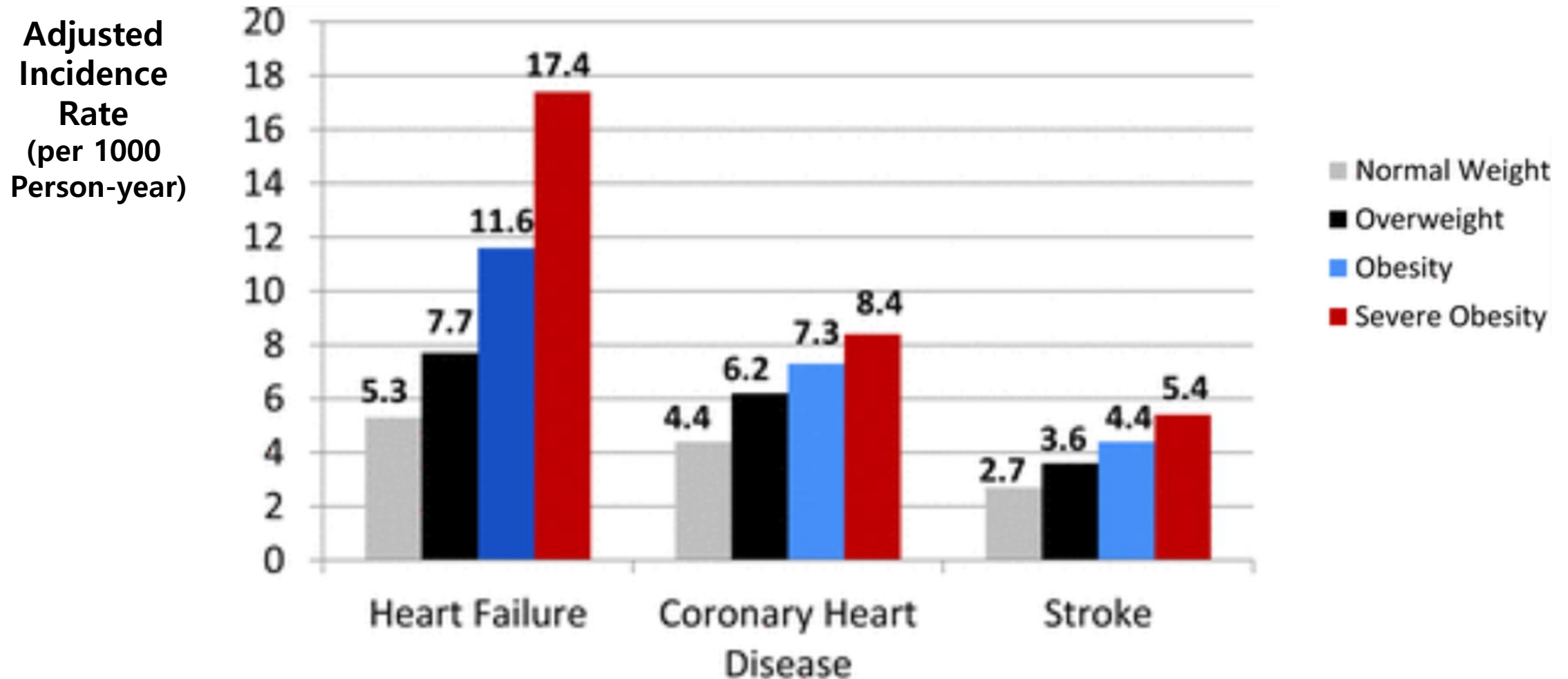
Adopted Quetelet index as
a nutritional indicator

for investigating the
relationship between diet
and cardiovascular
diseases in 1972

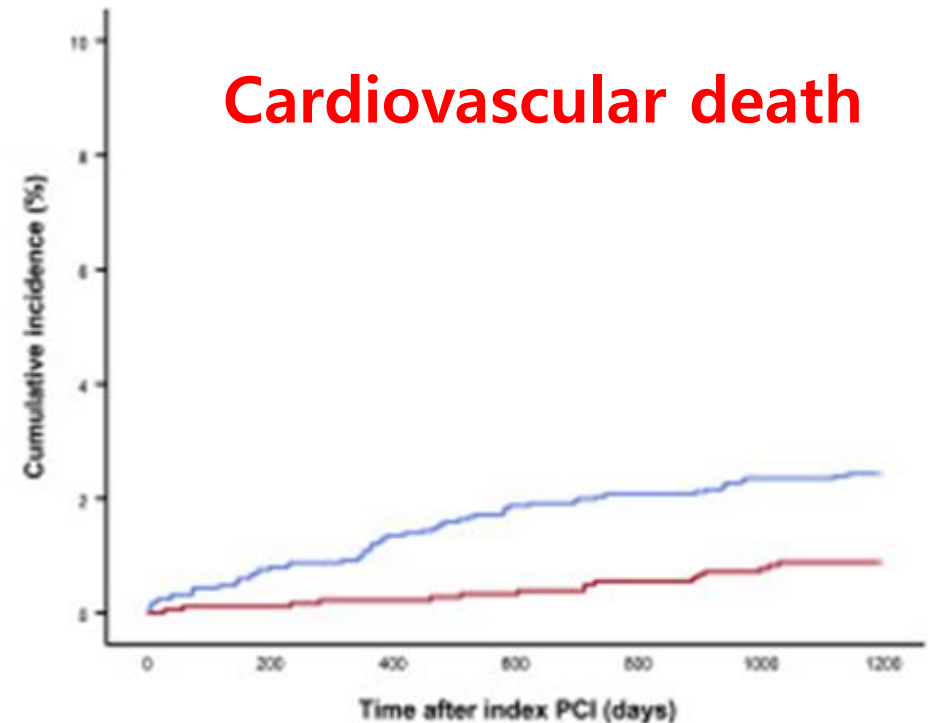
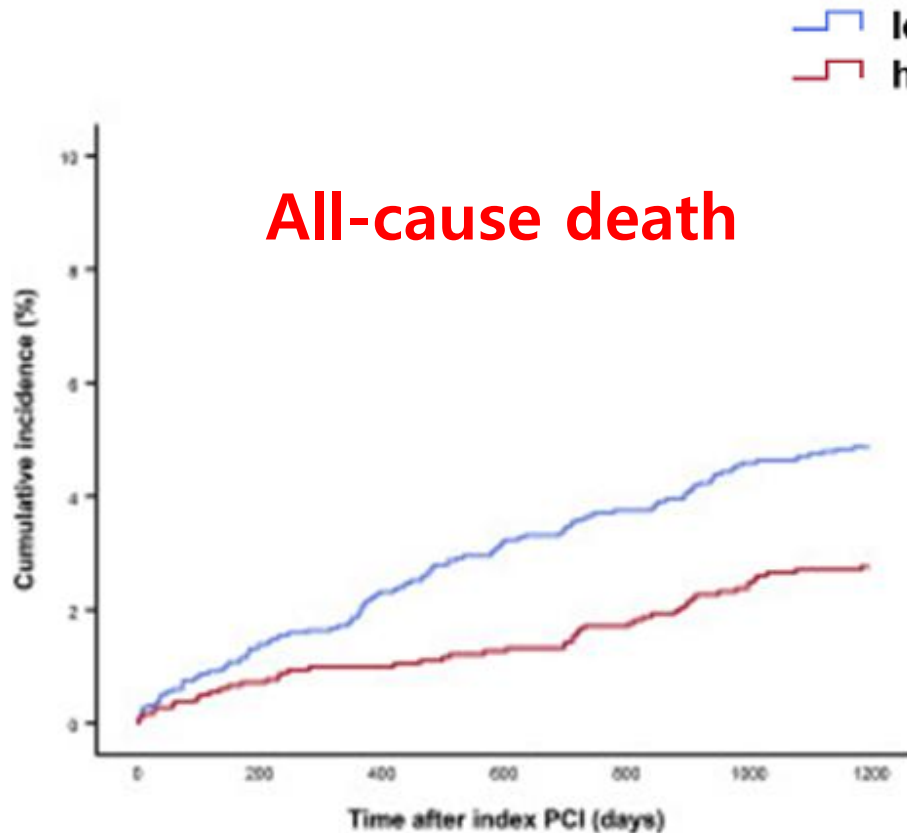


**Issues in Obesity,
originated from BMI**

Obesity Severity based on BMI and Incident Cardiovascular Disease

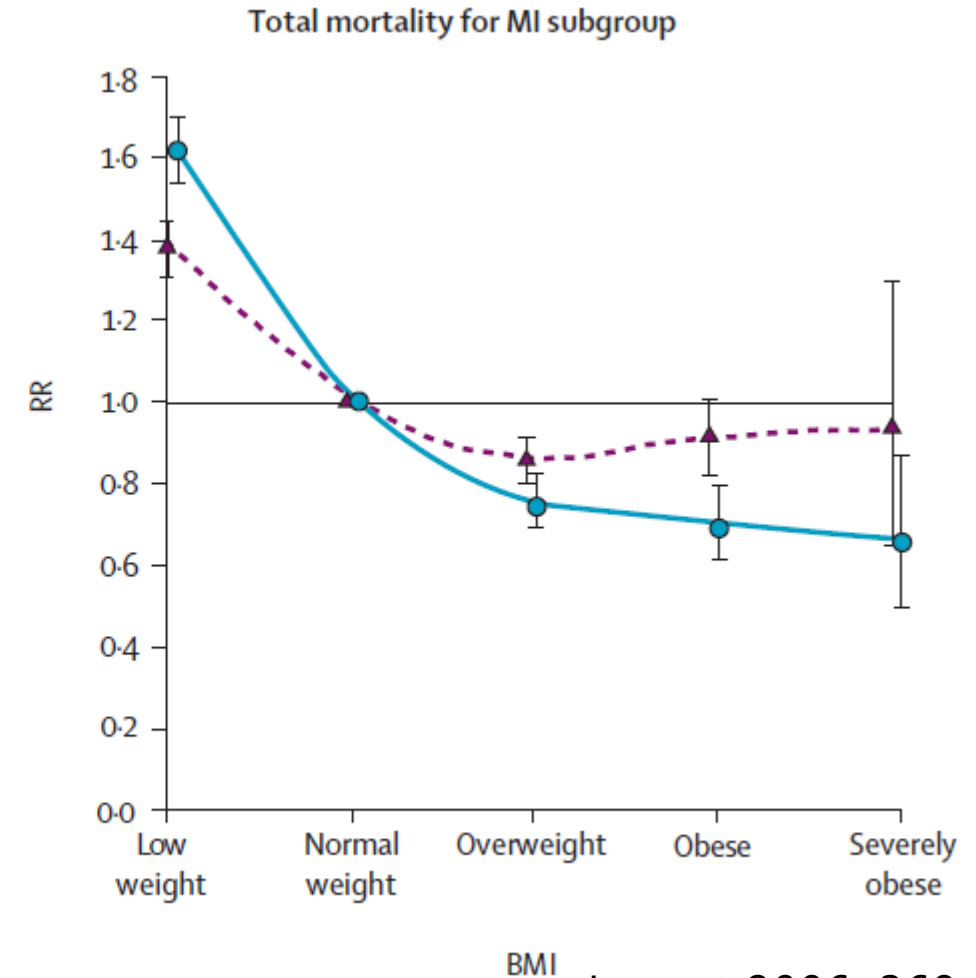
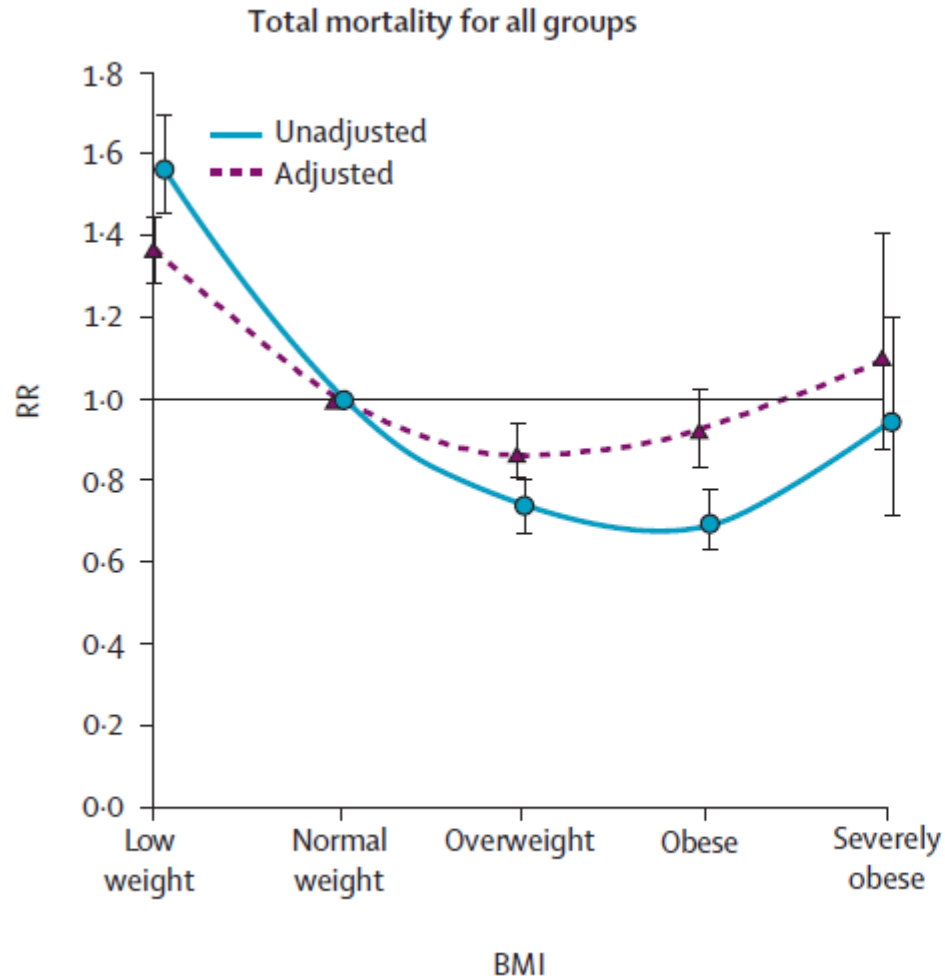


Outcomes comparing lower and higher BMI groups at three-year follow up of MI



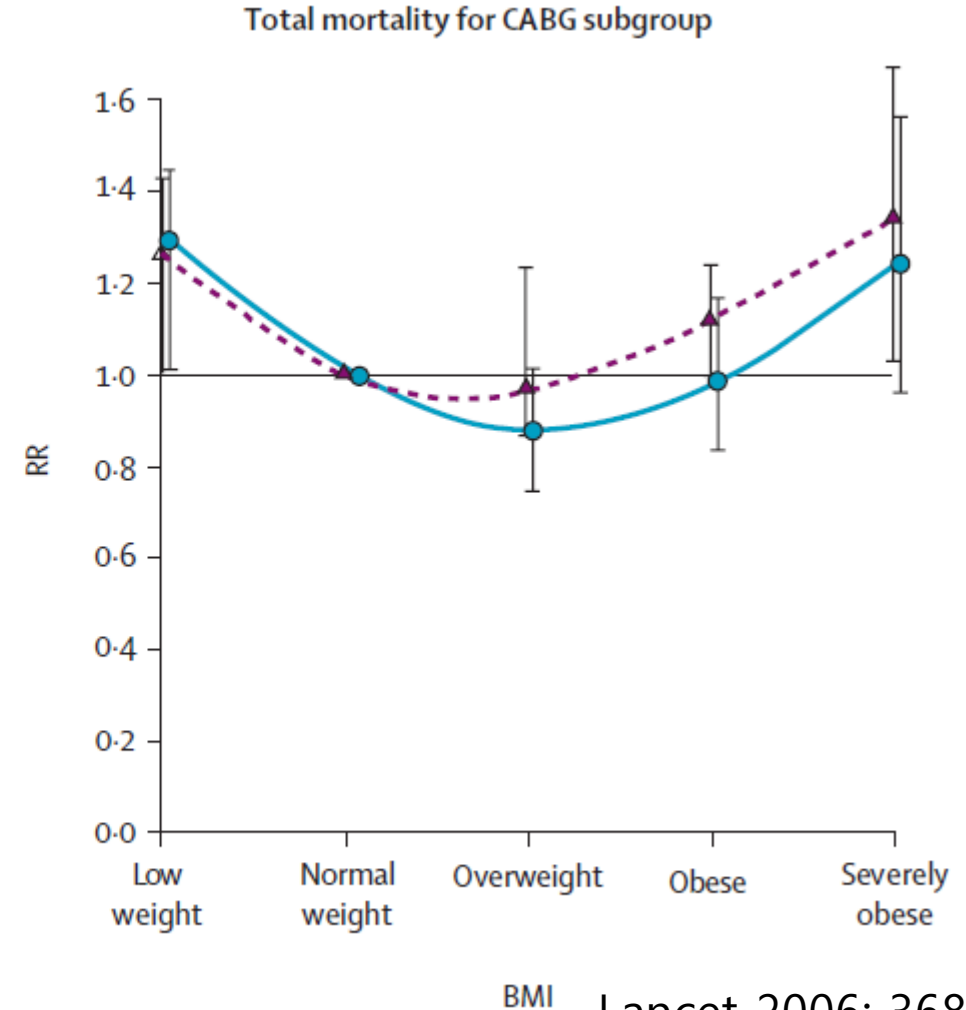
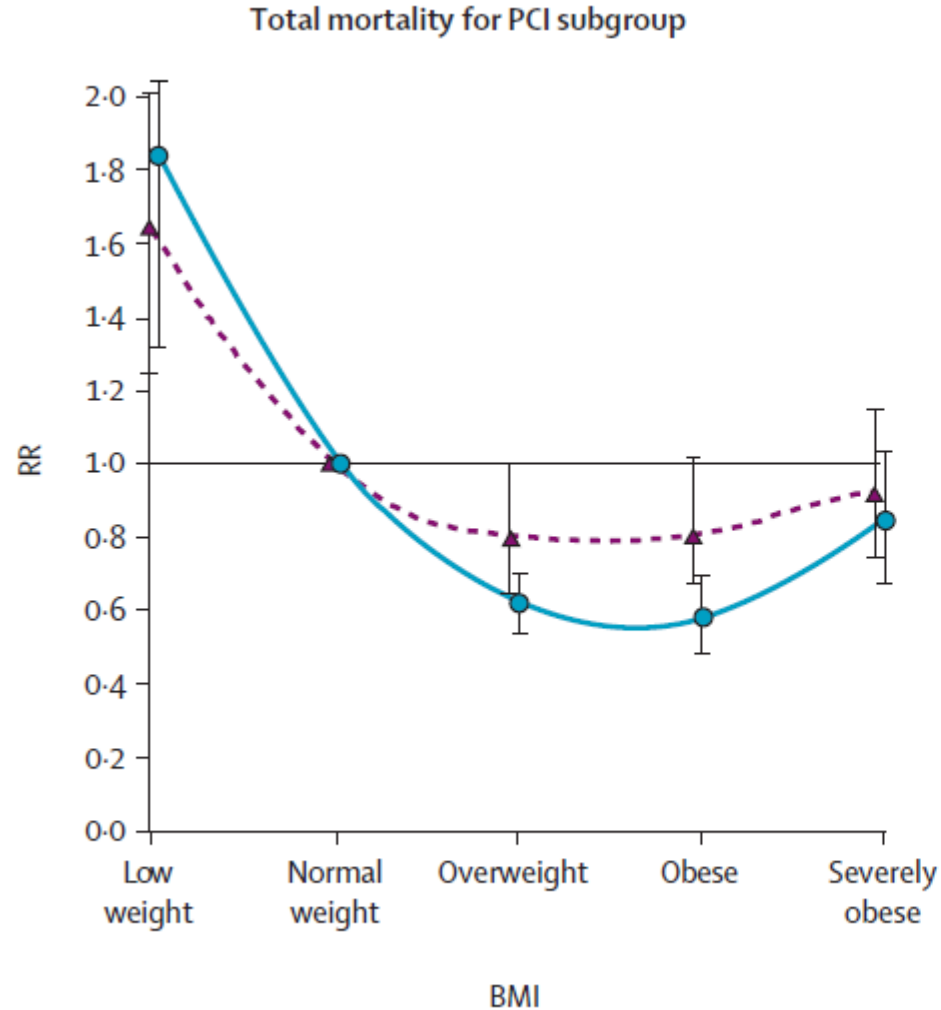
Relative Risk for Total mortality in patients with Coronary artery disease by BMI groups

Meta-analysis with 40 studies with 250,152 patients



RR for Total mortality in patients with CAD after PCI or CABG by BMI groups

Meta-analysis with 40 studies with 250,152 patients

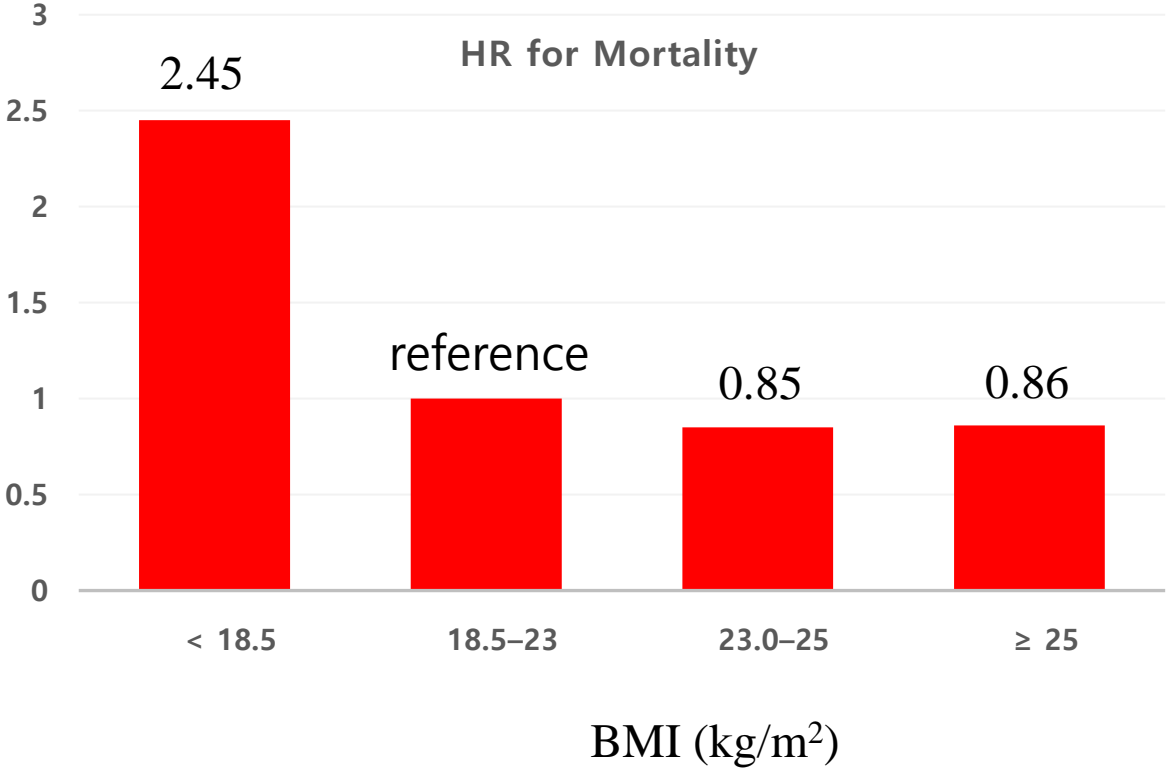


Adjusted Hazard Ratios for 10 year All-cause Death in Postmenopausal Korean Women

Korean National Health Insurance Service data



< 0.001,

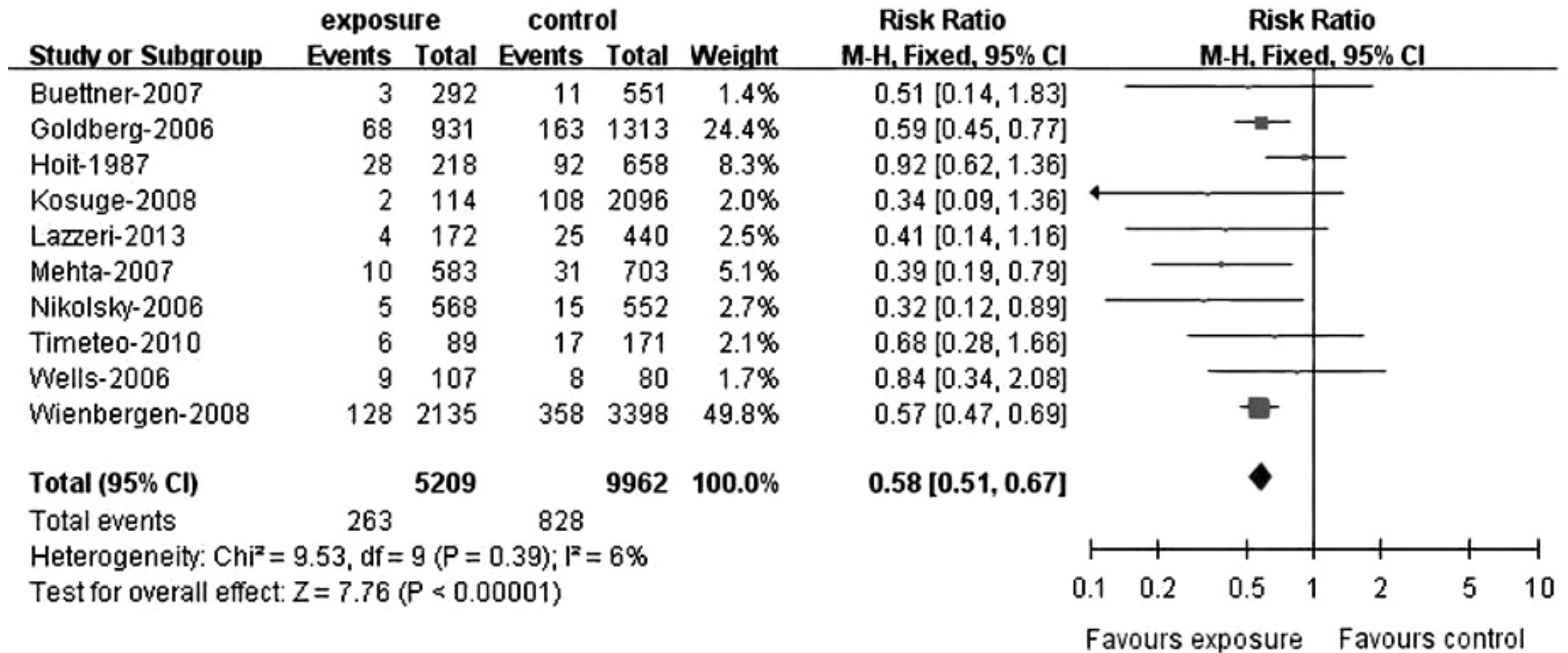


Kim BT, unpublished data

Relative Risk of mortality after acute MI according to BMI category

a meta-analysis of prospective studies

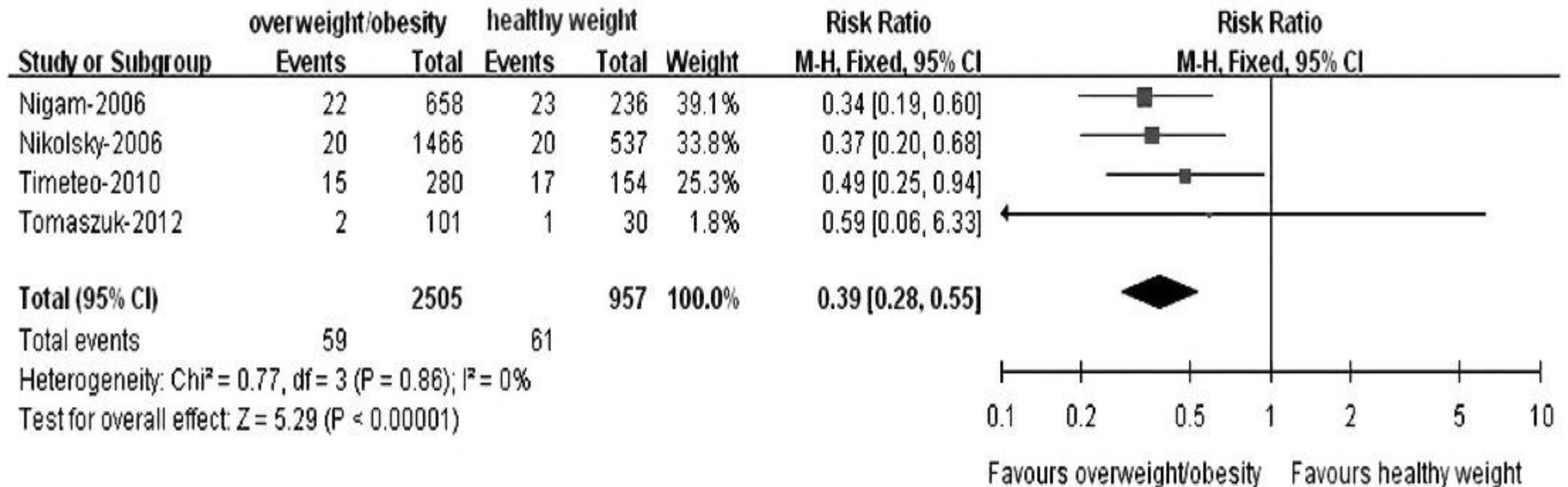
obesity(exposure) vs healthy weight(control)



Risk of short-term mortality after acute MI according to BMI category

a meta-analysis of prospective studies

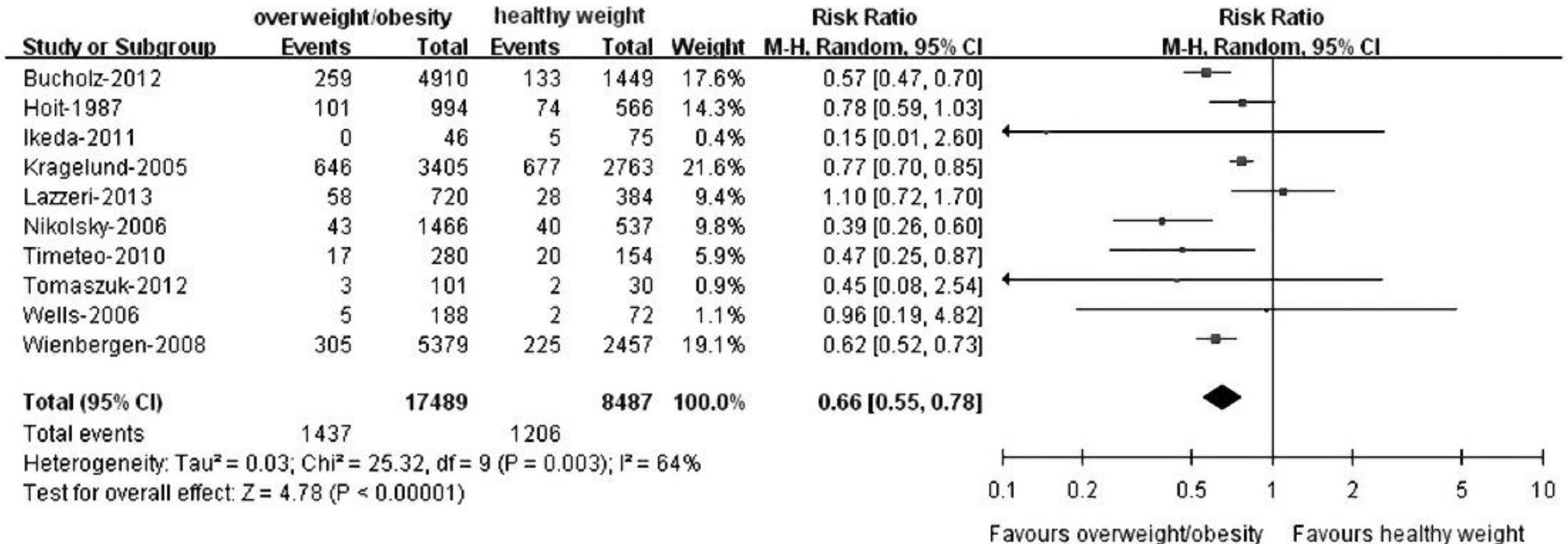
short-term mortality



Risk of medium-term mortality after acute MI according to BMI category

a meta-analysis of prospective studies

medium-term mortality



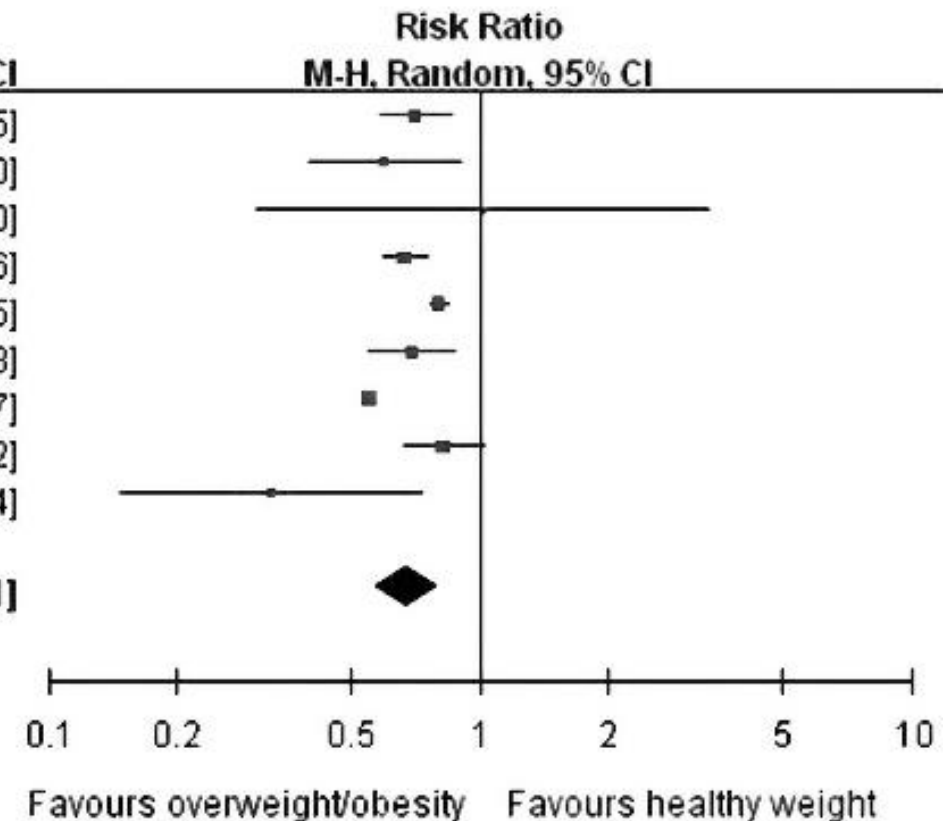
Risk of long-term mortality after acute MI according to BMI category

a meta-analysis of prospective studies

long-term mortality

Study or Subgroup	overweight/obesity		healthy weight		Weight	Risk Ratio		Risk Ratio	
	Events	Total	Events	Total		M-H, Random, 95% CI	M-H, Random, 95% CI		
Aronson-2010	244	1513	146	644	13.6%	0.71 [0.59, 0.85]			
Buettner-2007	50	1116	41	551	8.8%	0.60 [0.40, 0.90]			
Ikeda-2011	4	46	6	70	1.9%	1.01 [0.30, 3.40]			
Kennedy-2005	500	3482	408	1906	14.8%	0.67 [0.60, 0.76]			
Kragelund-2005	1430	3405	1438	2763	15.6%	0.81 [0.77, 0.85]			
Nigam-2006	154	658	79	236	12.6%	0.70 [0.56, 0.88]			
O'Brien-2014	6261	22043	6353	12422	15.8%	0.56 [0.54, 0.57]			
Rana-2004	198	1291	113	607	13.0%	0.82 [0.67, 1.02]			
Tomaszuk-2012	10	101	9	30	3.8%	0.33 [0.15, 0.74]			
Total (95% CI)		33655		19229	100.0%	0.68 [0.57, 0.81]			
Total events	8851		8593						

Heterogeneity: $\tau^2 = 0.05$; $\chi^2 = 169.15$, $df = 8$ ($P < 0.00001$); $I^2 = 95\%$
 Test for overall effect: $Z = 4.27$ ($P < 0.00001$)



Relative risks of Mortality with Body Mass Index among Patients with Chronic Obstruct Pulmonary Disease

1.1.2 Overweight vs Normal

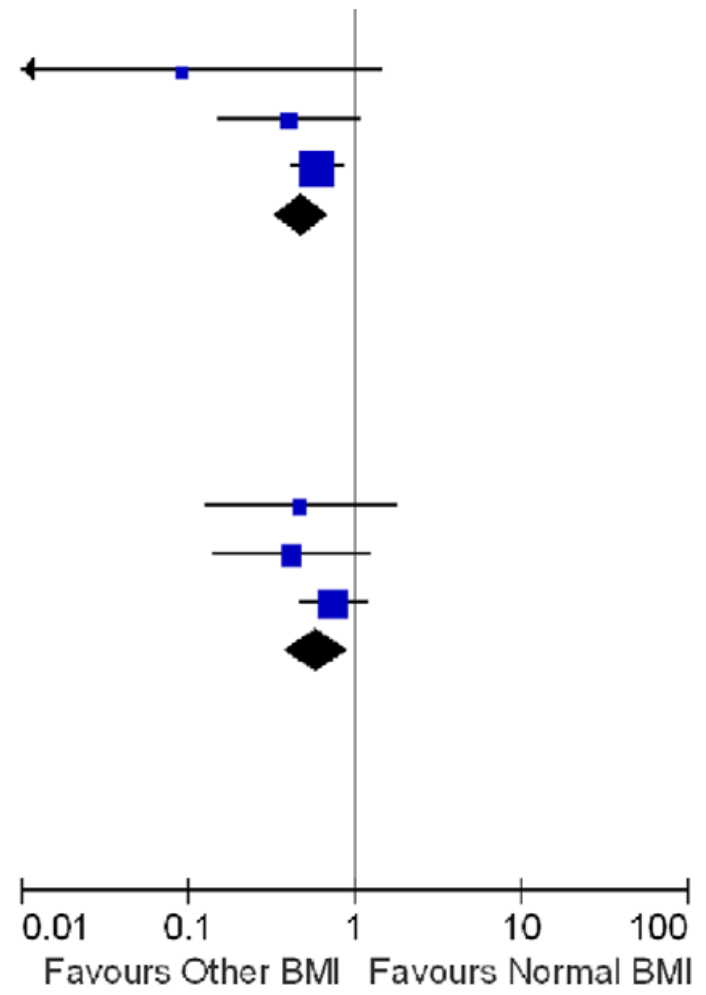
Chang et al, 2007	0	13	16	41	14.1%	0.09 [0.01, 1.42]
Gunen et al, 2005	5	120	15	144	23.2%	0.40 [0.15, 1.07]
Marti et al, 2006	21	47	38	50	62.7%	0.59 [0.41, 0.84]
Subtotal (95% CI)		180		235	100.0%	0.47 [0.33, 0.68]

Total events 26 69
 Heterogeneity: $\text{Chi}^2 = 2.91, \text{df} = 2 (P = 0.23); I^2 = 31\%$
 Test for overall effect: $Z = 4.00 (P < 0.0001)$

1.1.3 Obese vs Normal

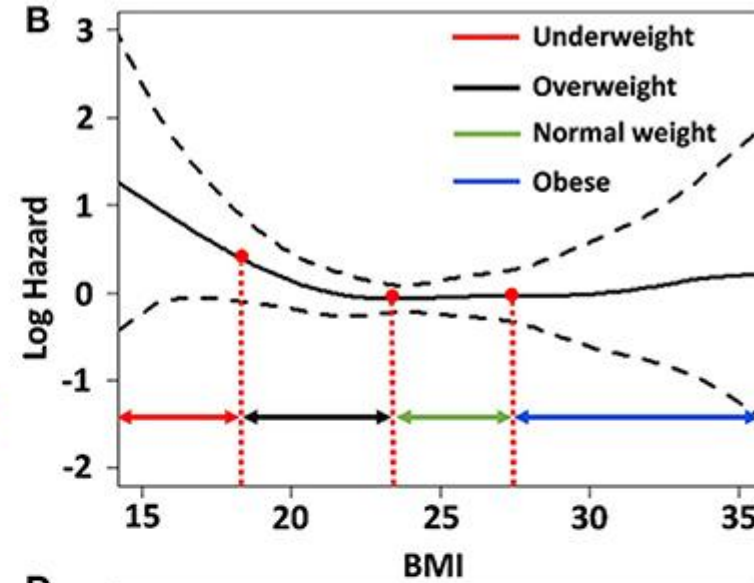
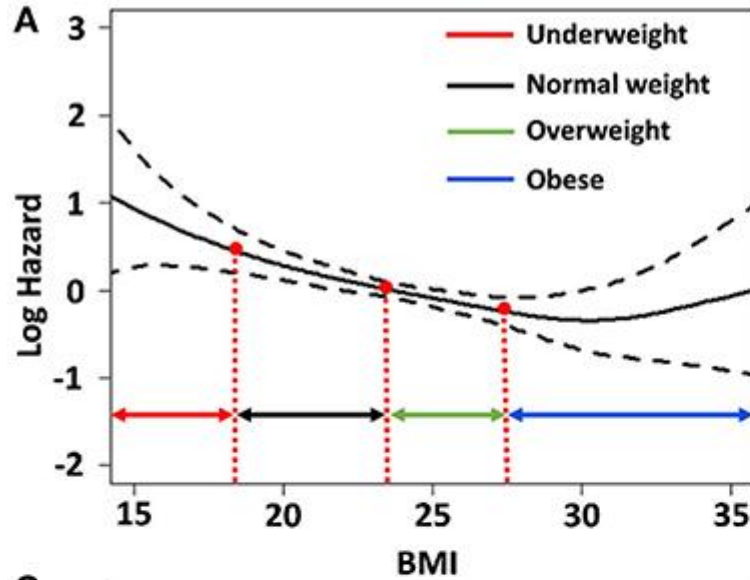
Chang et al, 2007	2	11	16	41	18.3%	0.47 [0.13, 1.73]
Collins et al, 2010	4	93	15	144	31.8%	0.41 [0.14, 1.21]
Marti et al, 2006	9	16	38	50	49.8%	0.74 [0.47, 1.17]
Subtotal (95% CI)		120		235	100.0%	0.59 [0.38, 0.91]

Total events 15 69
 Heterogeneity: $\text{Chi}^2 = 1.52, \text{df} = 2 (P = 0.47); I^2 = 0\%$
 Test for overall effect: $Z = 2.38 (P = 0.02)$



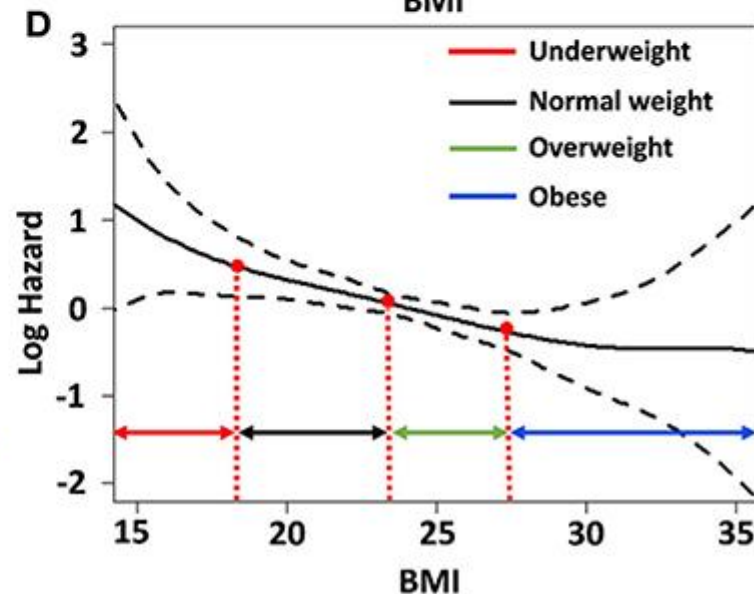
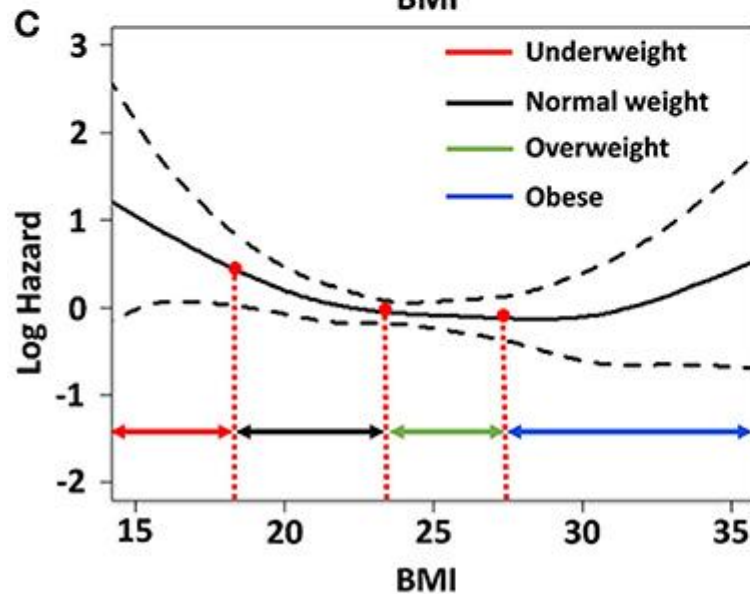
Relative hazards of body mass index (BMI) on CVD outcomes after adjustment

all-cause mortality



fatal stroke

fatal CV events



other-cause mortality

The definition of Obesity

- a " chronic, relapsing, multi-factorial, neurobehavioral disease, wherein an **increase in body fat** promotes **adipose tissue dysfunction** and **abnormal fat** mass physical forces, resulting in adverse metabolic, biomechanical, and psychosocial health consequences."

The Obesity Medicine Association,

3 Questions about BMI

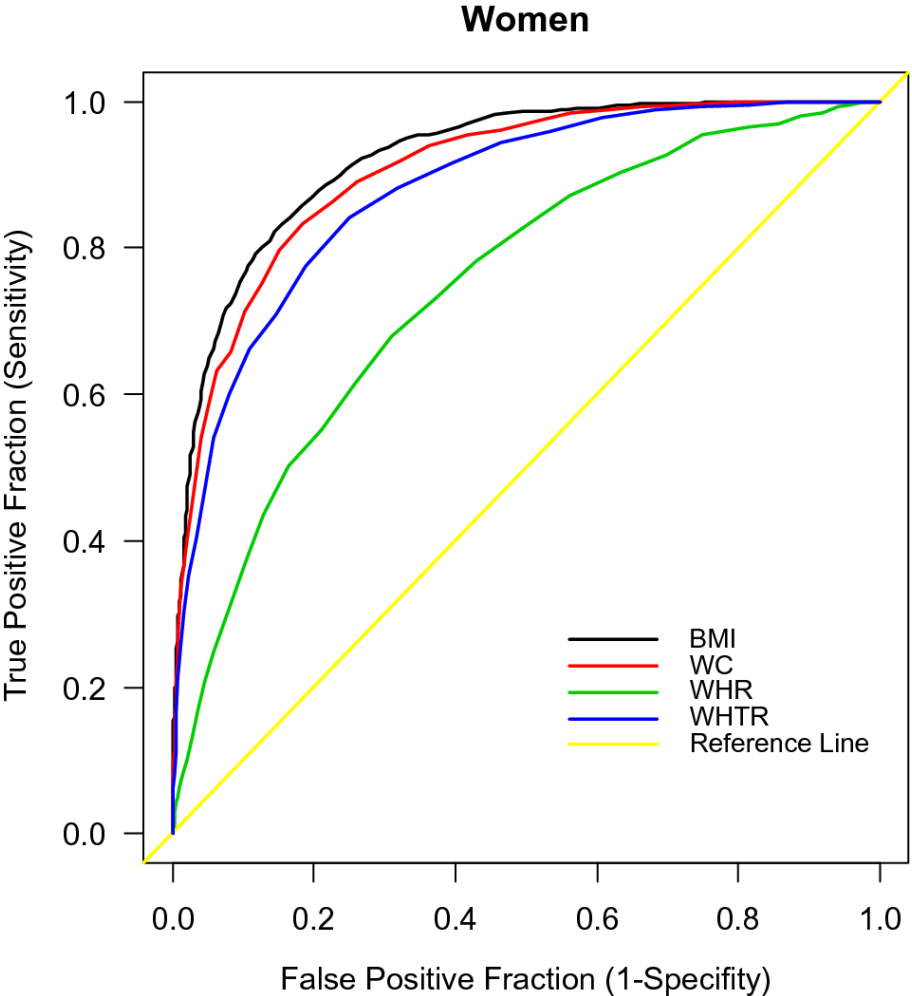
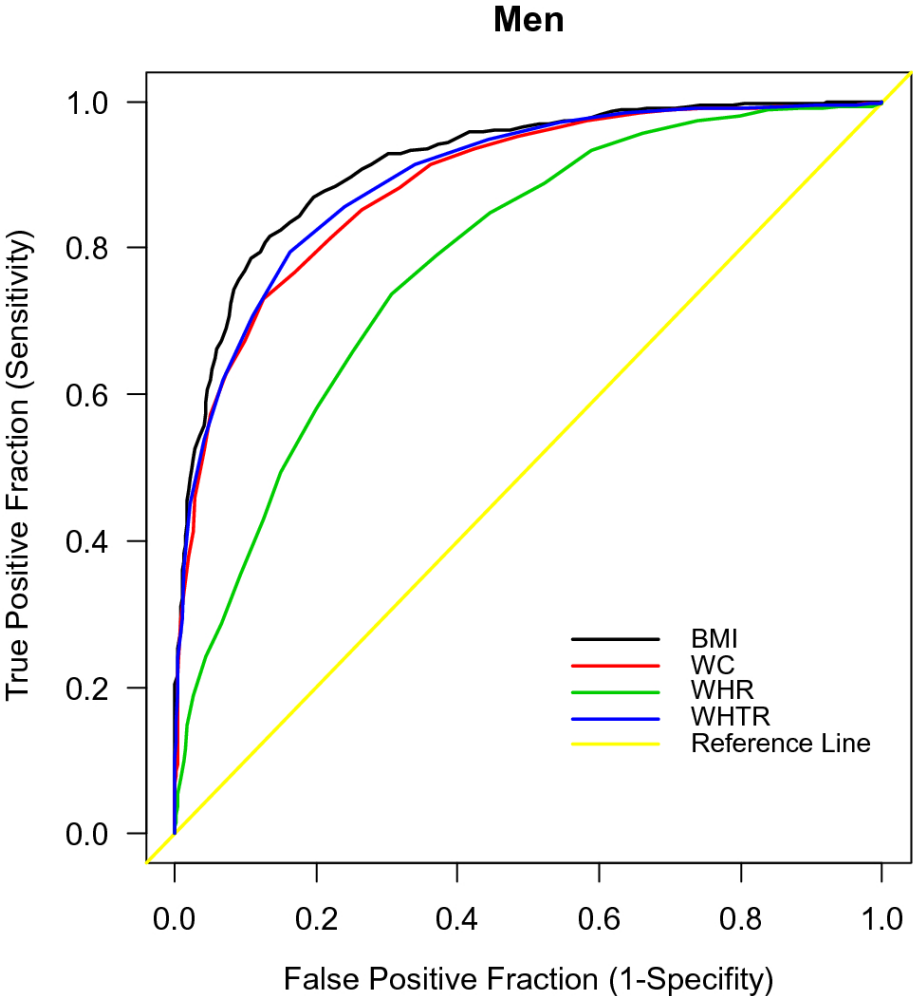
1. First, is BMI a good measure of excess adiposity in an individual patient ?
2. Second, does BMI provide information about body fat distribution, which is a more important predictor to the risk of obesity related consequences than the BMI itself?
3. Finally, does BMI provide any insights into the heterogeneity of obesity or its genetic, metabolic, physiological or psychological origins?

Correlations between body mass index/percent body fat and blood pressure, glucose and lipids.

	Body mass index		Percent body fat	
	Correlation coefficient	Partial correlation coefficient	Correlation coefficient	Partial correlation coefficient
SBP (mmHg)	0.403†	0.071†	0.098†	-0.006
DBP (mmHg)	0.409†	0.054†	0.075†	0.008
FPG (mM)	0.249†	0.02	0.088†	0.038*
TC (mM)	0.169†	-0.019	0.112†	0.078†
TG (mM)	0.350†	-0.021	0.049†	0.051†
HDL-C (mM)	-0.432†	-0.063†	-0.003	-0.050†
LDL-C (mM)	0.189†	0.016	0.097†	0.079†

*P < 0.05 and †P < 0.01 (t-test). Confounding factors such as age, gender, lifestyle, and family history were adjusted in the partial correlation analysis

ROC curves for BMI, WC, WHR, and WHTR in screening for cardiovascular risk factors



Cut-off values for anthropometric indicators of obesity, based on optimal Body Fat % cut-offs for screening cardiovascular risk factors

	AUC (95% CI)	Optimal cut-off	Sensitivity (%)	Specificity (%)	Youden (%)	DLR (+)	DLR (-)	P
Men								
BMI	0.915 (0.902, 0.929)	28.1	81.6	86.6	68.1	6.1	0.2	0.000001
WC	0.887 (0.871, 0.903)	100.0	73.0	87.3	60.4	5.8	0.3	0.000001
WHR	0.781 (0.759, 0.803)	0.96	73.8	69.4	43.2	2.4	0.4	0.000001
WHTR	0.895 (0.880, 0.910)	0.57	79.5	83.6	63.1	4.9	0.2	0.000001
Women								
BMI	0.922 (0.913, 0.931)	27.5	83.3	84.5	67.8	5.4	0.2	0.000001
WC	0.905 (0.895, 0.915)	87.0	83.3	81.6	64.9	4.5	0.2	0.000001
WHR	0.747 (0.730, 0.764)	0.85	67.9	69.0	36.9	2.2	0.5	0.000001
WHTR	0.878 (0.866, 0.889)	0.54	84.1	75.1	59.1	3.4	0.2	0.000001

BF% in newly estimated non-obesity and obesity categories based on BMI, WC, WHR, WHTR, stratified by gender.

	Men		Women		
	BF < 25.8%	BF ≥ 25.8%	BF < 37.1%	BF ≥ 37.1%	
BMI < 28.1	677 (82.0)	149 (18.0)	BMI < 27.5	1522 (87.1)	225 (12.9)
BMI ≥ 28.1	105 (13.7)	659 (86.3)	BMI ≥ 27.5	279 (20.0)	1,119 (80.0)
WC < 100.0	683 (75.8)	218 (24.2)	WC < 87.0	1,470 (86.7)	225 (13.3)
WC ≥ 100.0	99 (14.4)	590 (85.6)	WC ≥ 87.0	331 (22.8)	1,119 (77.2)
WHR < 0.96	543 (71.9)	212 (28.1)	WHR < 0.85	1,287 (73.3)	470 (26.8)
WHR ≥ 0.96	239 (28.6)	596 (71.4)	WHR ≥ 0.85	514 (37.0)	874 (63.0)
WHTR < 0.57	654 (79.8)	166 (20.2)	WHTR < 0.54	1,352 (86.3)	214 (13.7)
WHTR ≥ 0.57	128 (16.6)	642 (83.4)	WHTR ≥ 0.54	449 (28.4)	1,130 (71.6)

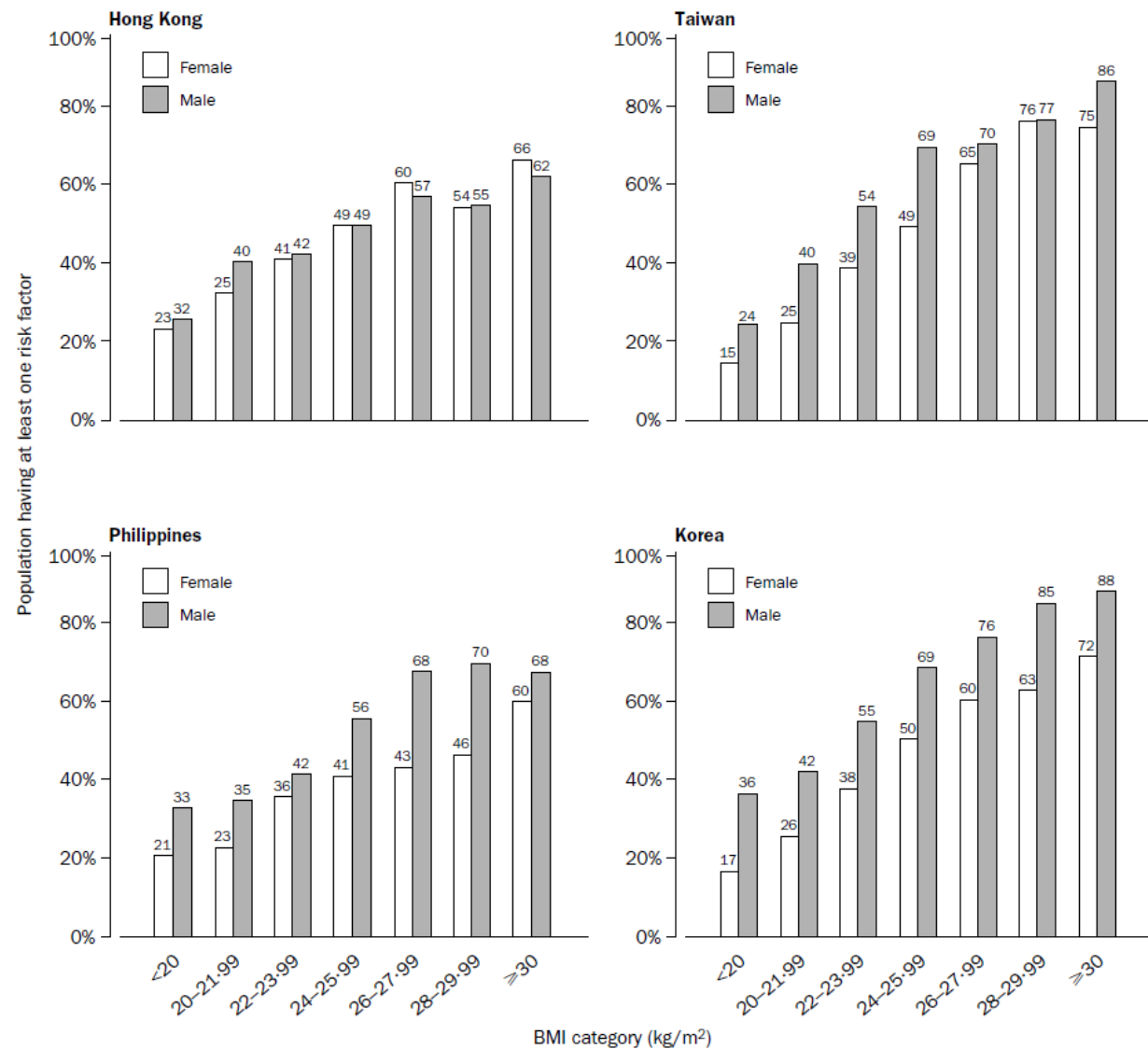
Odds ratios (and 95% CIs) for metabolic disorders using prediction models with BMI, WC, or both BMI and WC

the 3rd National Health and Nutrition Examination Survey.

	BMI alone	WC alone	BMI and WC	
			BMI	WC
Men (<i>n</i> = 7385)				
Hypertension	1.13 (1.11, 1.15) ²	1.04 (1.01, 1.05) ²	1.06 (1.00, 1.12) ²	1.03 (1.01, 1.05) ²
Hypercholesterolemia	1.07 (1.04, 1.10) ²	1.03 (1.02, 1.04) ²	1.02 (0.97, 1.08)	1.02 (1.00, 1.04) ³
High LDL cholesterol	1.16 (1.12, 1.90) ²	1.02 (1.01, 1.14) ²	1.03 (0.94, 1.13)	1.01 (0.99, 1.04)
Low HDL cholesterol	1.20 (1.17, 1.23) ²	1.06 (1.05, 1.07) ²	1.03 (0.98, 1.09)	1.05 (1.03, 1.07) ²
High triacylglycerol	1.20 (1.17, 1.23) ²	1.07 (1.06, 1.08) ²	1.05 (0.99, 1.12)	1.05 (1.03, 1.08) ²
Metabolic syndrome	1.20 (1.15, 1.25) ²	1.07 (1.06, 1.08) ²	1.07 (0.99, 1.18)	1.04 (1.02, 1.07) ²
Women (<i>n</i> = 7539)				
Hypertension	1.13 (1.10, 1.15) ²	1.05 (1.04, 1.06) ²	1.02 (0.97, 1.07)	1.05 (1.03, 1.06) ²
Hypercholesterolemia	1.08 (1.06, 1.10) ²	1.03 (1.03, 1.04) ²	1.00 (0.97, 1.04)	1.03 (1.02, 1.05) ²
High LDL cholesterol	1.08 (1.04, 1.11) ²	1.04 (1.02, 1.05) ²	0.97 (0.92, 1.02)	1.05 (1.02, 1.07) ²
Low HDL cholesterol	1.11 (1.08, 1.13) ²	1.04 (1.02, 1.05) ²	1.03 (0.99, 1.08)	1.03 (1.01, 1.05) ²
High triacylglycerol	1.13 (1.11, 1.15) ²	1.06 (1.05, 1.07) ²	0.96 (0.92, 1.00)	1.08 (1.08, 1.09) ²
Metabolic syndrome	1.15 (1.12, 1.19) ²	1.06 (1.05, 1.08) ²	1.01 (0.95, 1.07)	1.06 (1.04, 1.09) ²

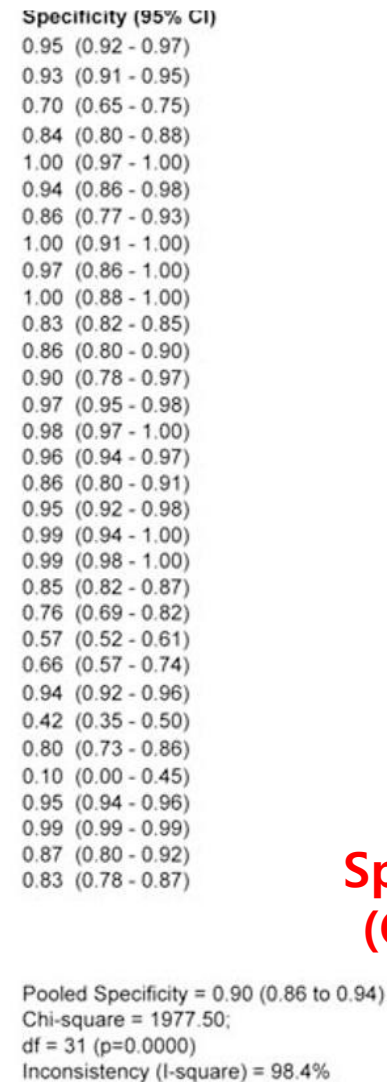
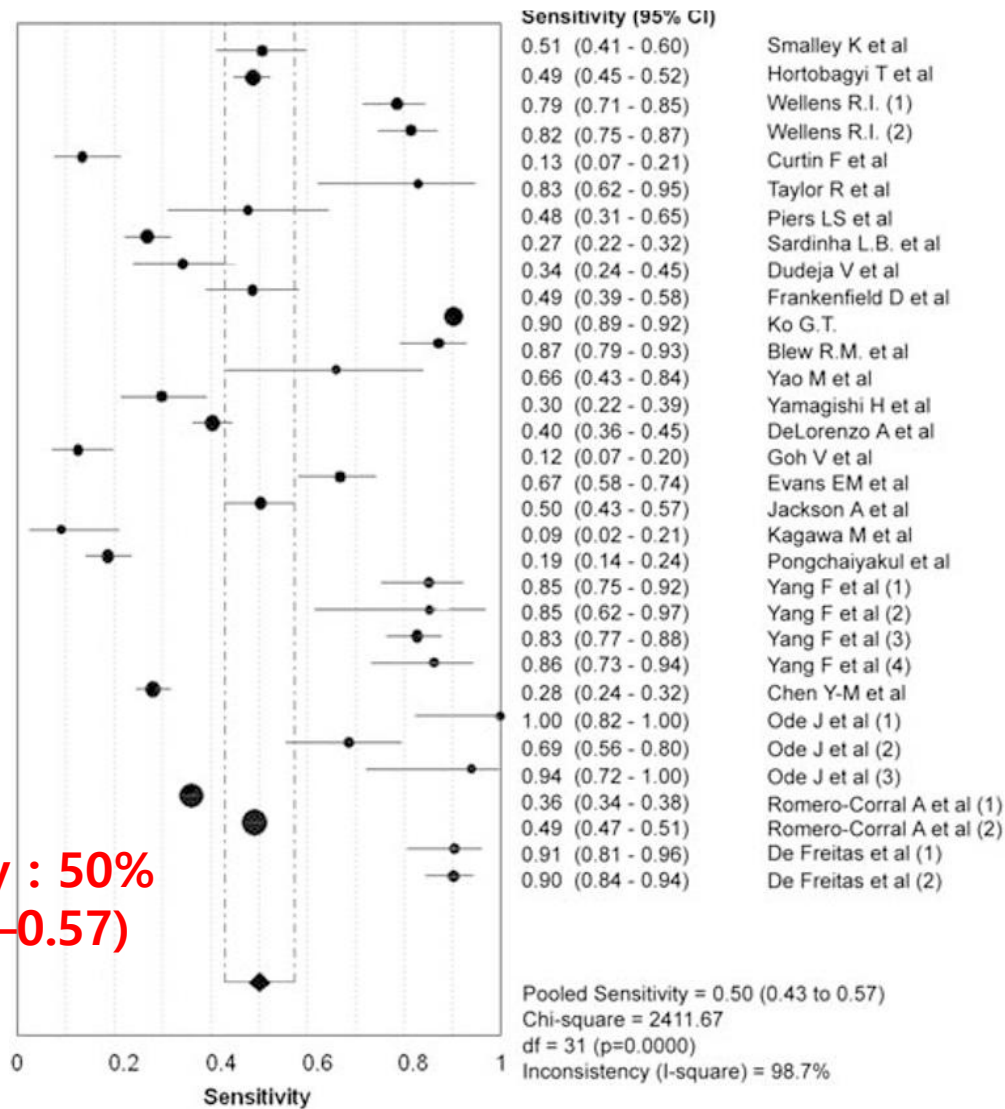
The ORs were adjusted for age, race, physical activity, smoking, alcohol intake, and the poverty-to-income ratio

Proportion of population in various body-mass index (BMI) categories with at least one risk factor for cardiovascular disease in Asians



Pooled sensitivity and specificity of BMI to identify obesity in Meta-analysis

Sensitivity : 50%
(CI: 0.43–0.57)



Specificity : 90%
(CI: 0.86–0.94).

BMI and Variance in body fat

Multiple regression analysis of body fat percentage versus body mass index, age, sex, and ethnicity for the total study population, New York City, 1986-1992

Body mass index	Regression coefficients				SEE	r^2 †
	Age	Sex‡	Ethnic group§	Intercept		
Mean ± SE†	Mean ± SE	Mean ± SE	Mean ± SE	Mean ± SE		
1.34 ± 0.07**				-6.07 ± 2.12*	8.48	0.26
1.23 ± 0.08**	0.15 ± 0.02**			-10.77 ± 2.07**	8.04	0.34
1.46 ± 0.06**	0.12 ± 0.01**	-11.61 ± 0.44**		-10.02 ± 1.46**	5.68	0.67
1.47 ± 0.06**	0.12 ± 0.01**	-11.61 ± 0.44**	-0.22 ± 0.49	-10.13 ± 1.48**	5.68	0.67

* $p < 0.01$; ** $p < 0.0001$.

† r^2 , explained variance of the model; SE, standard error; SEE, standard error of the estimate.

‡ 0 = female; 1 = male.

§ 0 = white, 1 = black.

Body mass index alone accounted for only **25%** of the variance in body fat in men and in women

Uses and limitations of the BMI

Measurement	Value Rating
Estimation of body weight	****
Tracking population weight	****
Estimation of body fat	**
Estimating distribution of fat	0
Use in genetic studies	**
Pathophysiology of obesity	0
Phenotyping obesity	0

0 = not useful information; ** = some useful information; **** = valuable.

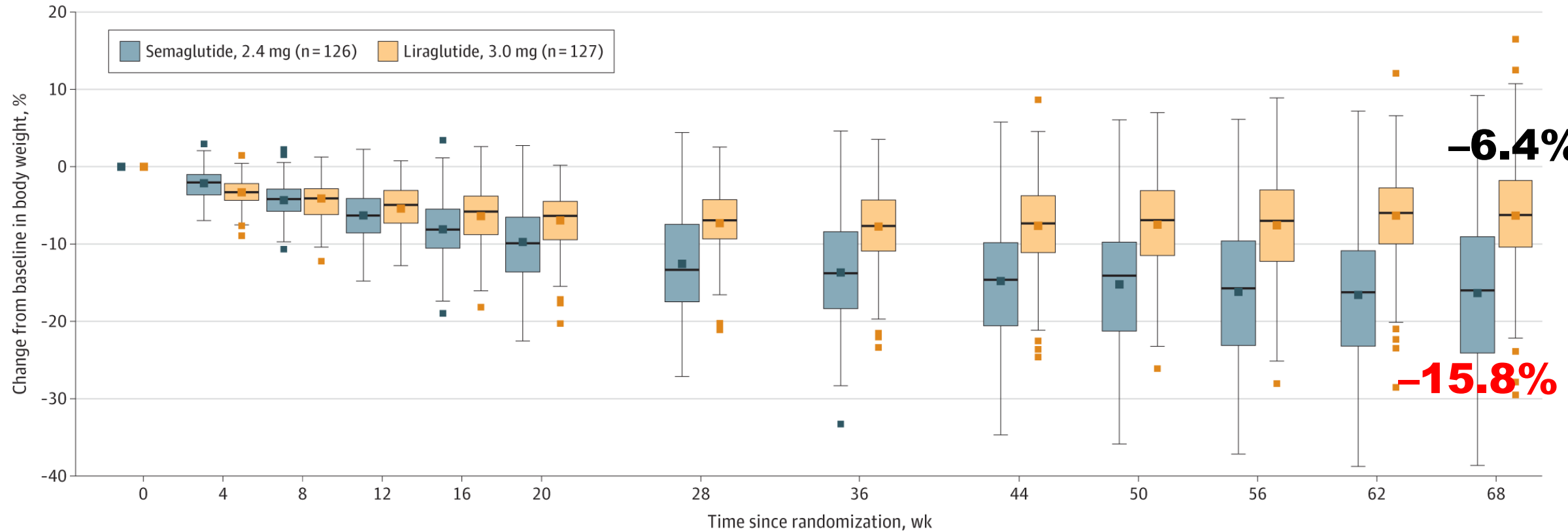
3 issues originated from BMI

- 1. The BMI has only a fair not a precise correlation with body fat mass.**
- 2. BMI does not provide any indication of how body fat is distributed.**
- 3. the BMI tells us nothing about the genetic, metabolic, physiological or psychological factors involved in the development of obesity.**

Obesities, beyond BMI



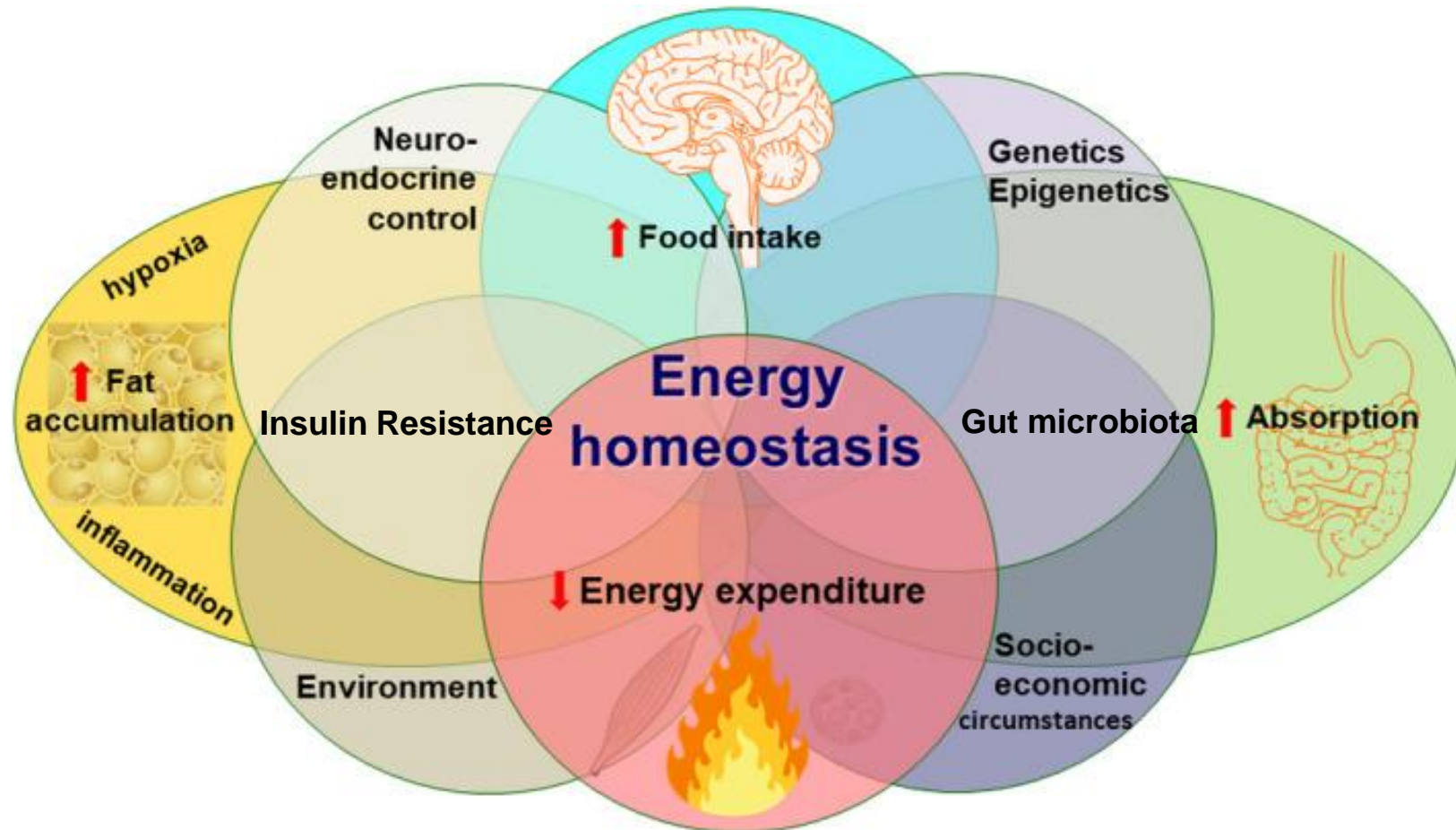
STEP 8 – Body Weight change (% on ITT)



No. of participants
 Semaglutide, 2.4 mg
 Liraglutide, 3.0 mg

	126	125	122	124	117	102	76	105	114	107	107	92	117
	127	124	124	125	118	101	66	102	99	98	110	88	117

Complex pathophysiology of Excessive Adiposity



4 pillars of Obesity Management

Lifestyle change
Diet & Exercise

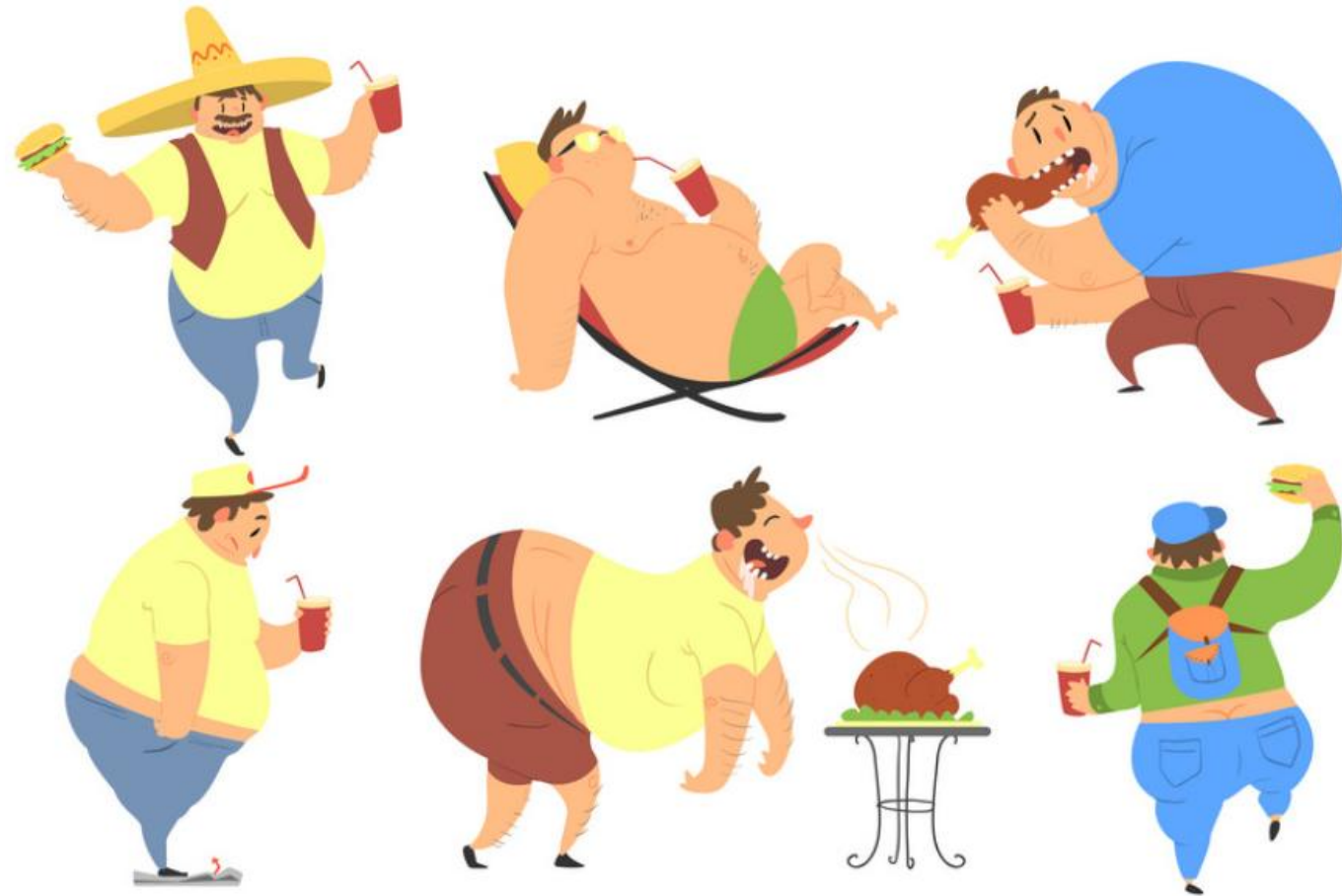
Behavioral
Interventions
: ~5% weight loss



Pharmacotherapy
: ~5–15% weight loss

Surgical
Interventions
: ~12–30% weight loss

Variety in phenotypes of Weight Gain



4 pillars of Obesity Management

Lifestyle change
Diet & Exercise

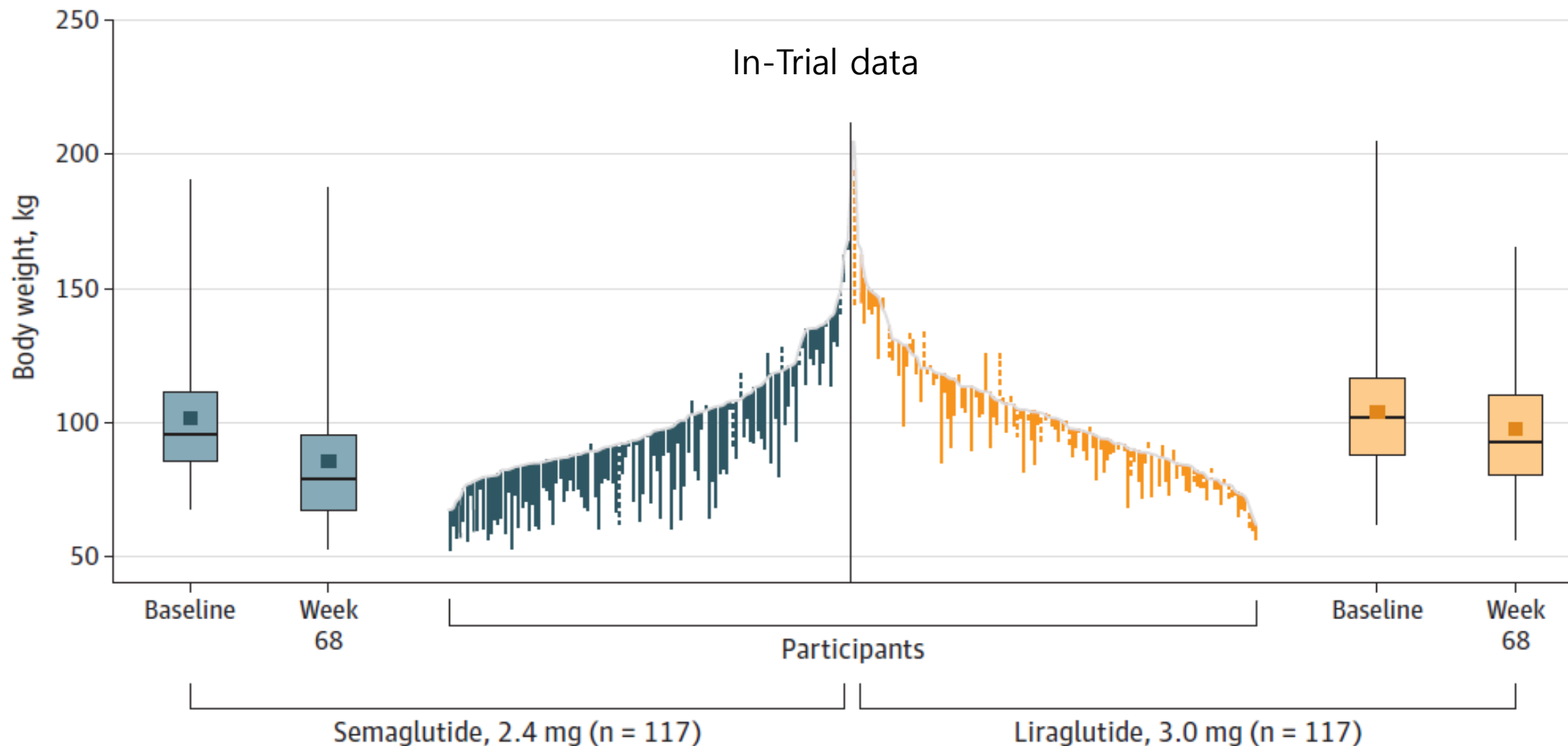
Behavioral
Interventions
: ~5% weight loss



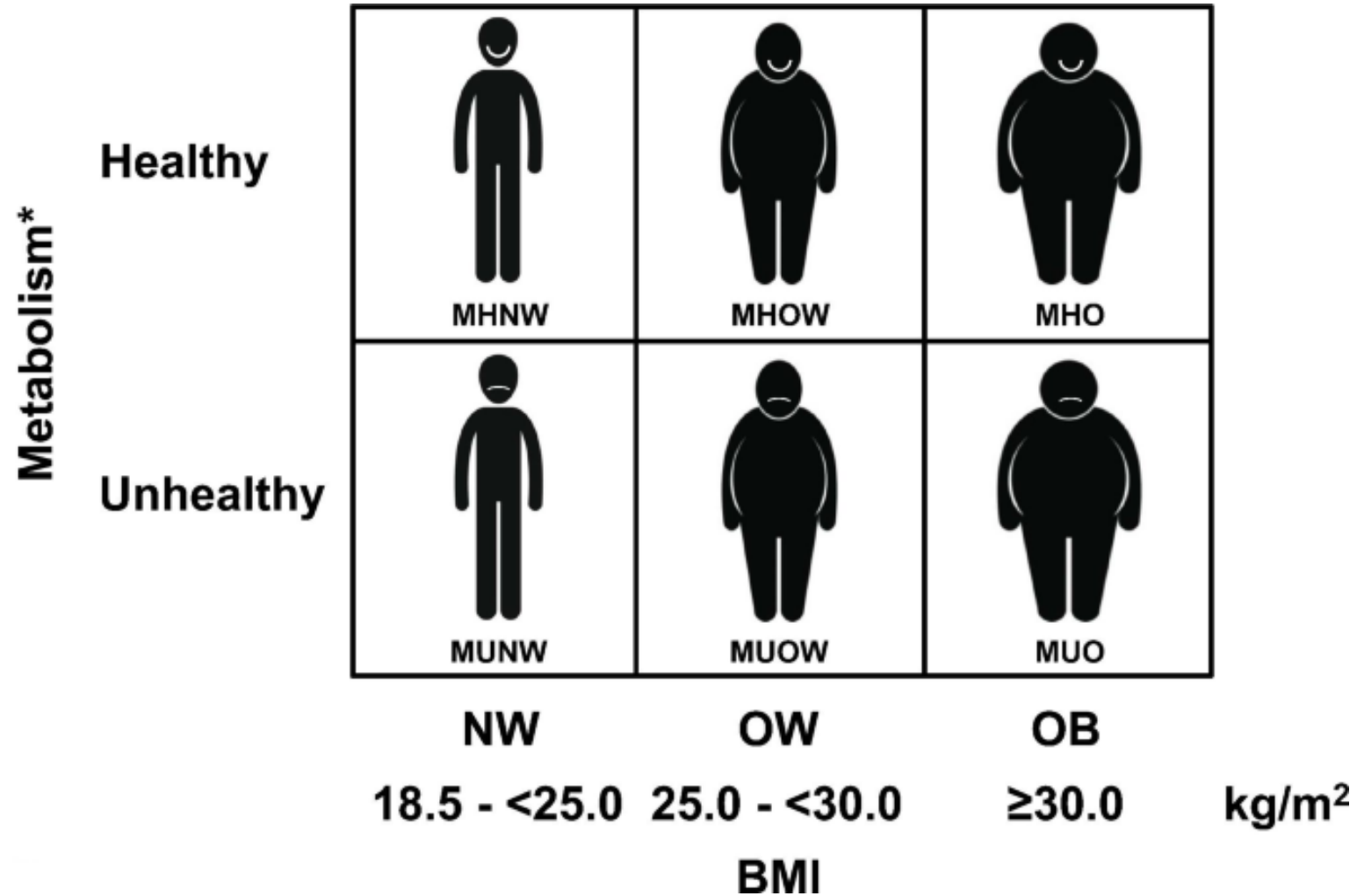
Pharmacotherapy
: ~5–15% weight loss

Surgical
Interventions
: ~12–30% weight loss

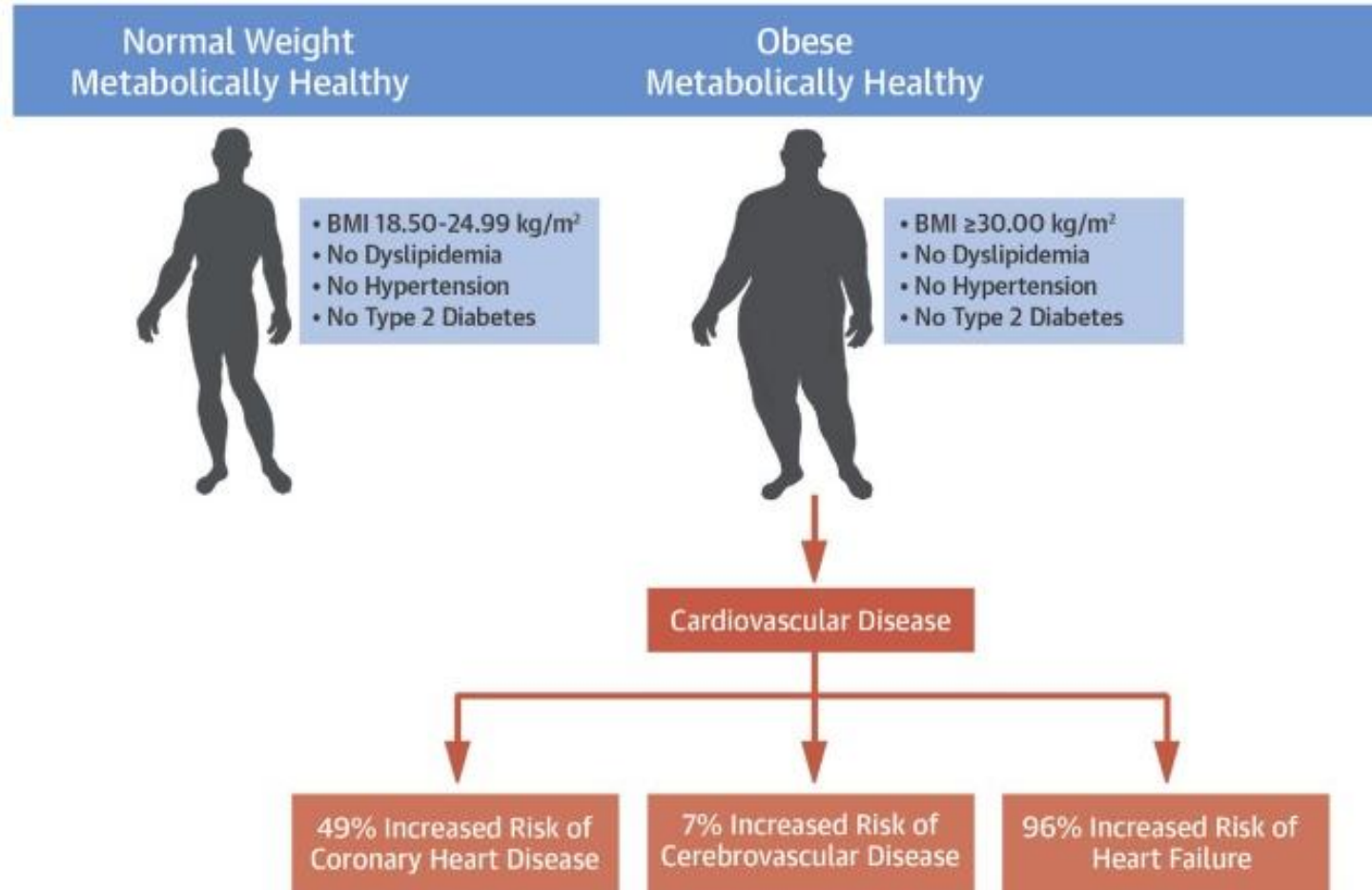
Change in Absolute Body Weight From Baseline to Week 68 for Individual Participants : STEP 8 trial



Metabolic phenotype of Obesity

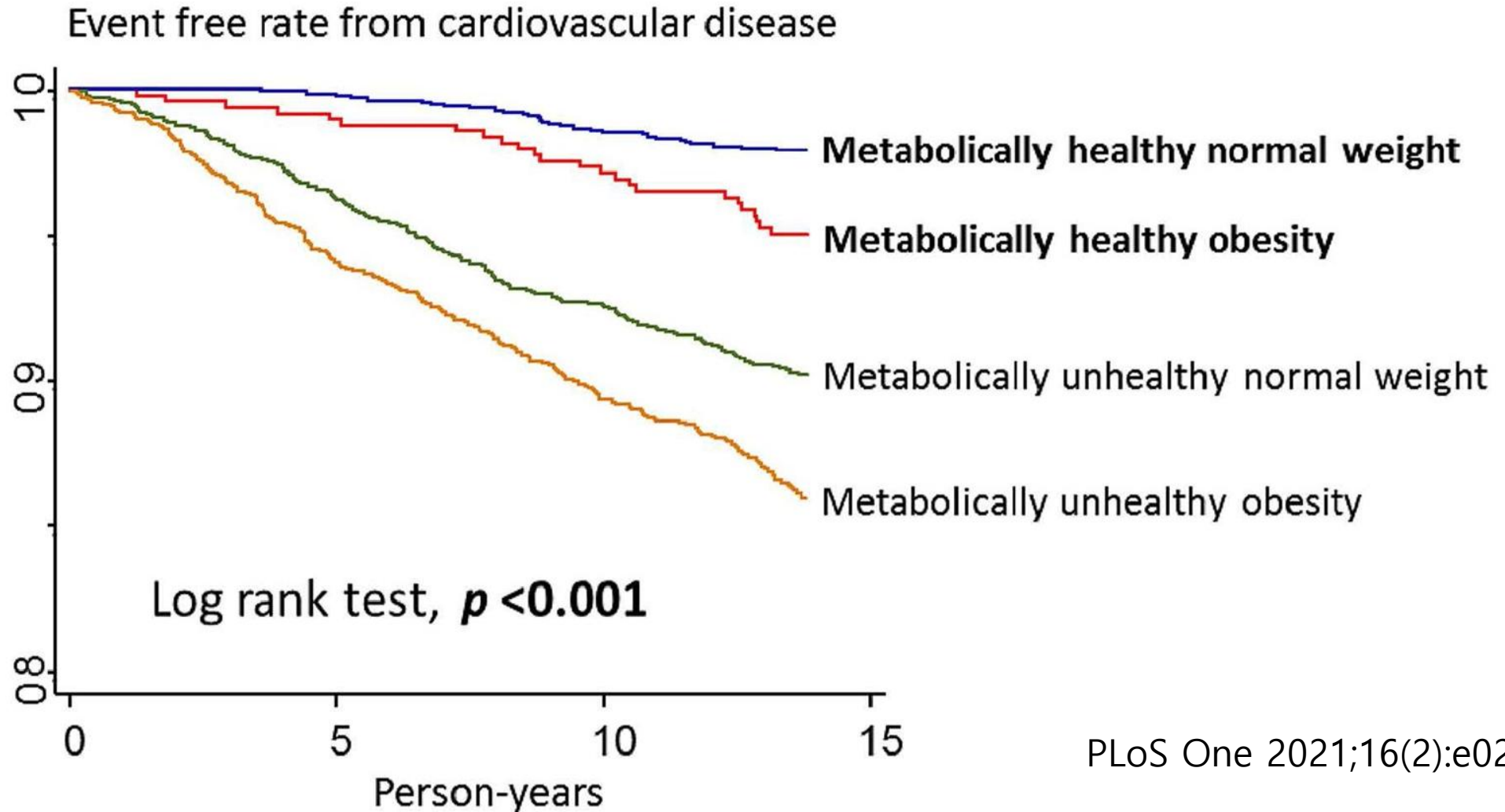


Metabolically Healthy Obese and Incident Cardiovascular Disease Events

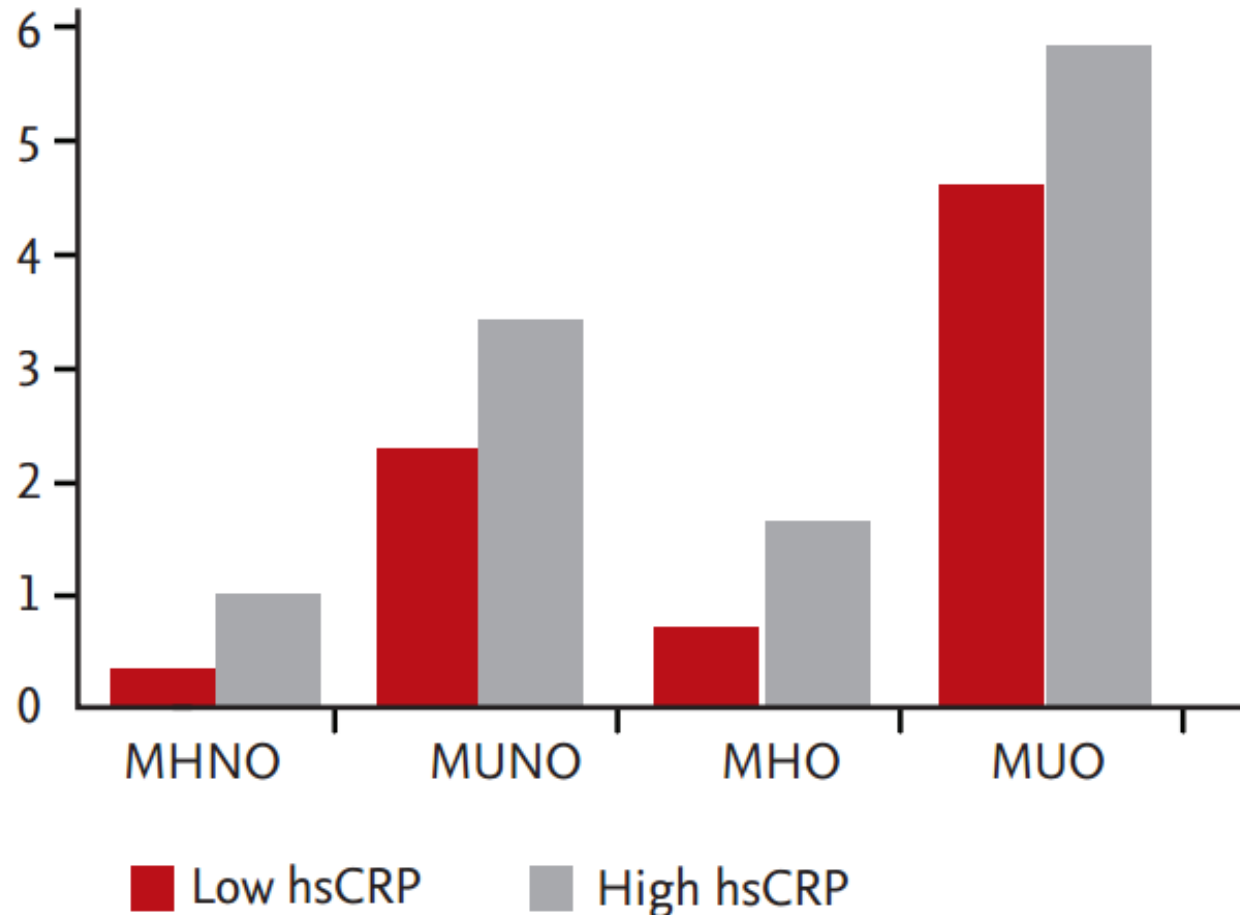


Association between metabolically healthy obesity/overweight and cardiovascular disease risk

A representative cohort study in Taiwan

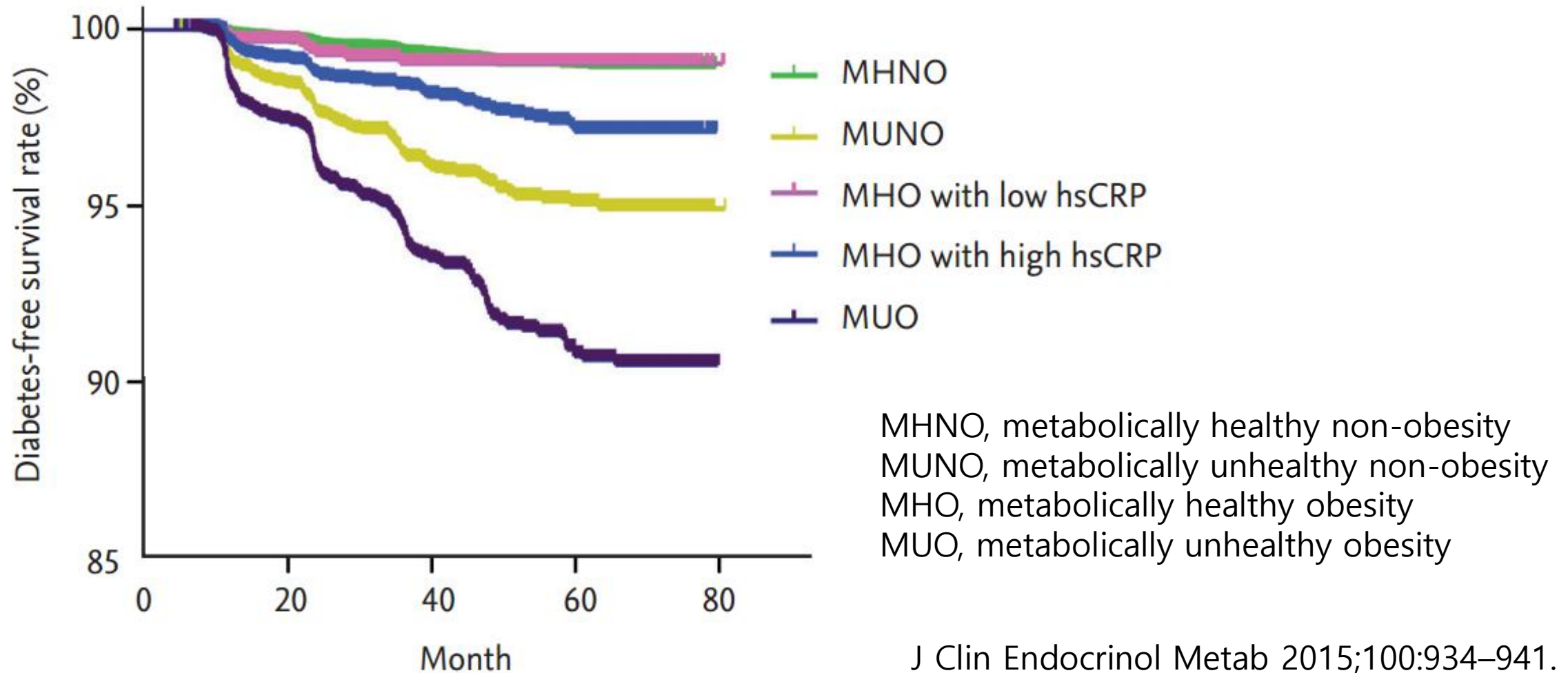


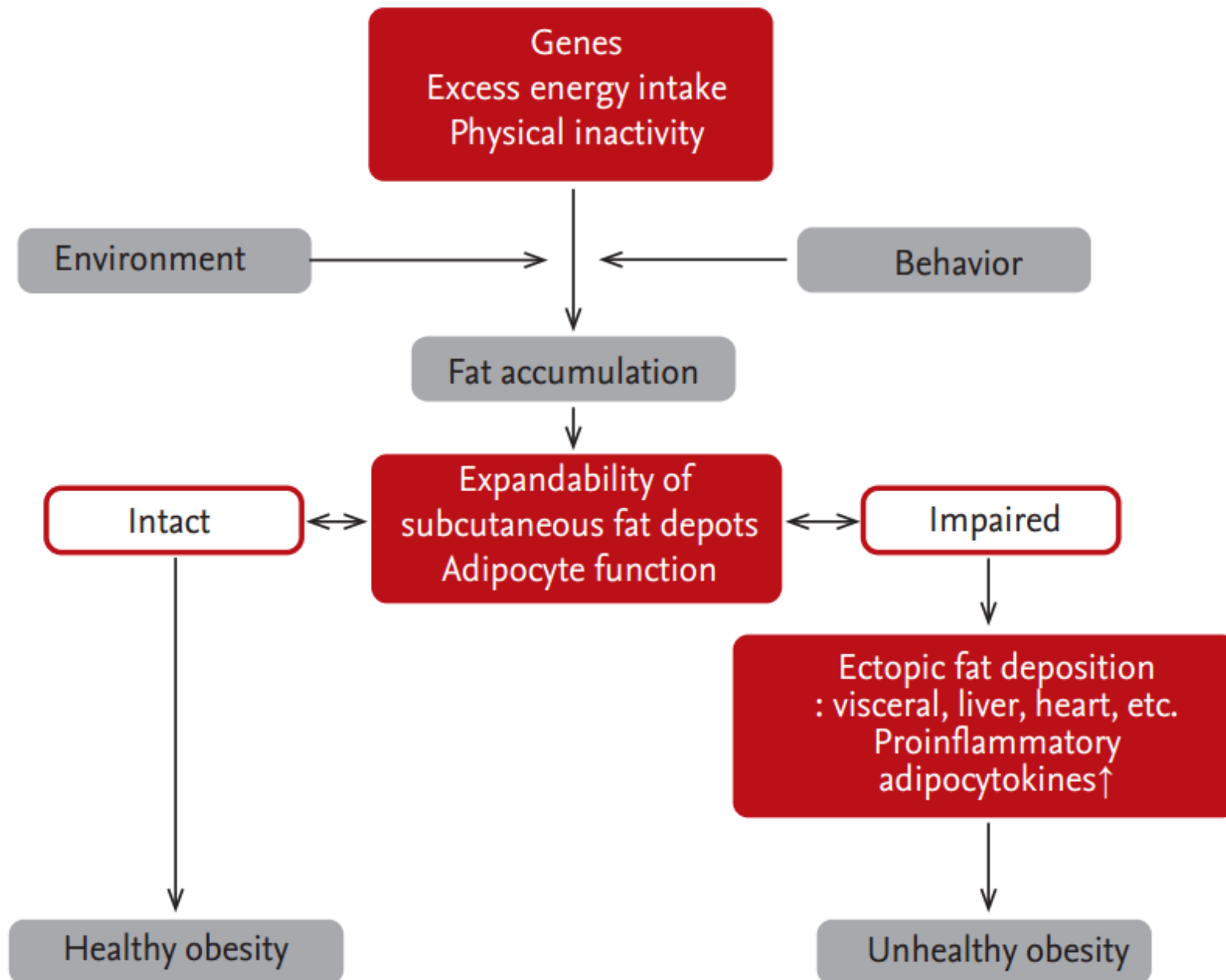
Interaction of low grade inflammation and obesity in the development of type 2 DM



MHNO, metabolically healthy non-obesity
MUNO, metabolically unhealthy non-obesity
MHO, metabolically healthy obesity
MUO, metabolically unhealthy obesity

Interaction of low grade inflammation and obesity in the development of type 2 DM



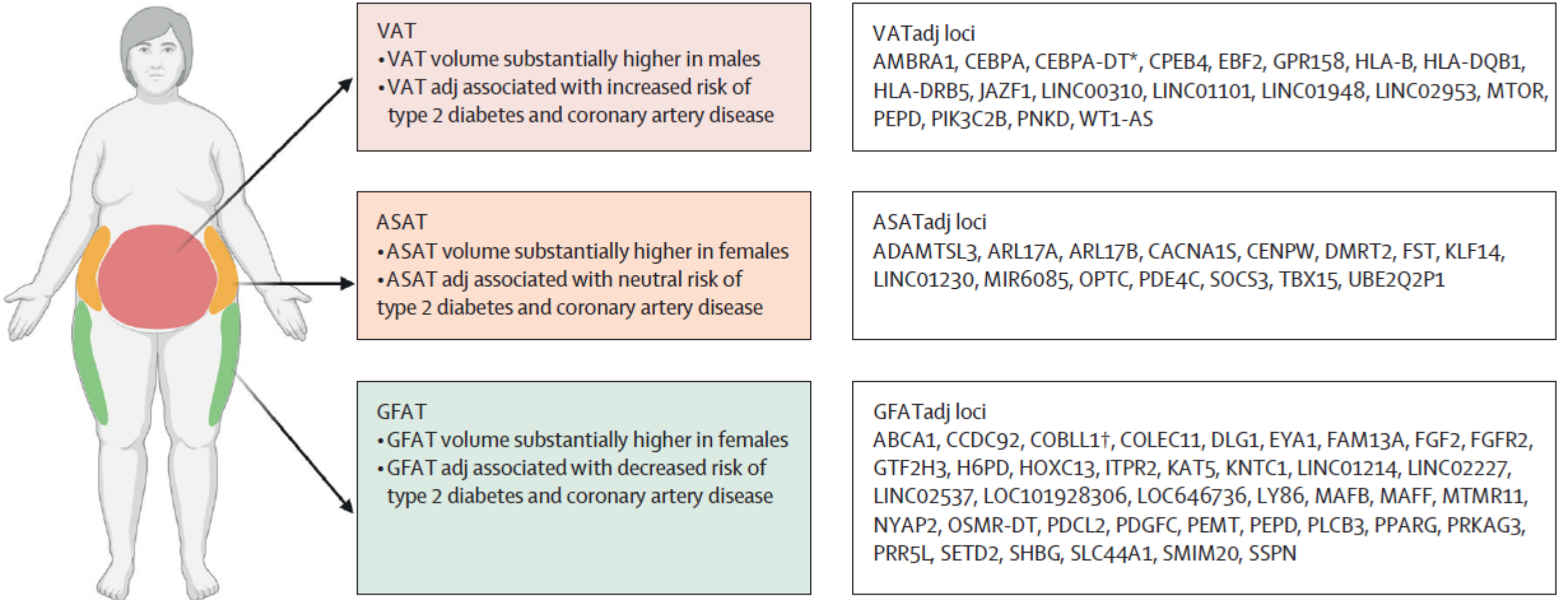


Genetic phenotype of Obesity

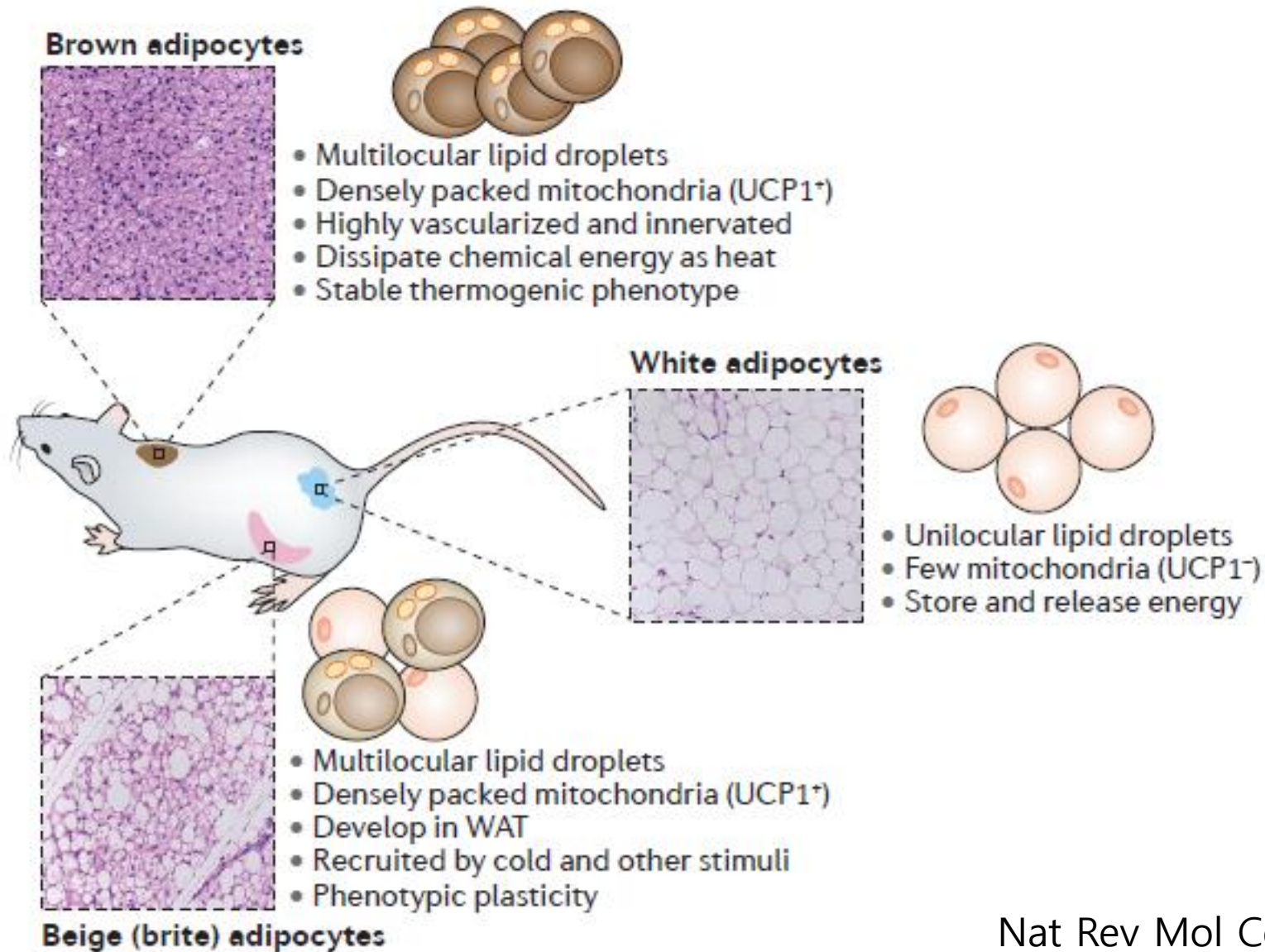
Association of selected GLP-1R polymorphisms with response to liraglutide

Polymorphism	Genotype	Poor responders <i>N</i> =37 <i>N</i> (%)	Strong responders <i>N</i> =20 <i>N</i> (%)	OR (95 % CI)	<i>P</i>
rs10305420	CC	12 (48.0)	13 (52.0)	0.27 (0.09–0.85)	0.025
	CT+TT	24 (77.4)	7 (22.6)		
rs6923761	GG	21 (77.8)	6 (22.2)	3.06 (0.96–9.74)	0.058
	GA+AA	16 (53.3)	14 (46.7)		

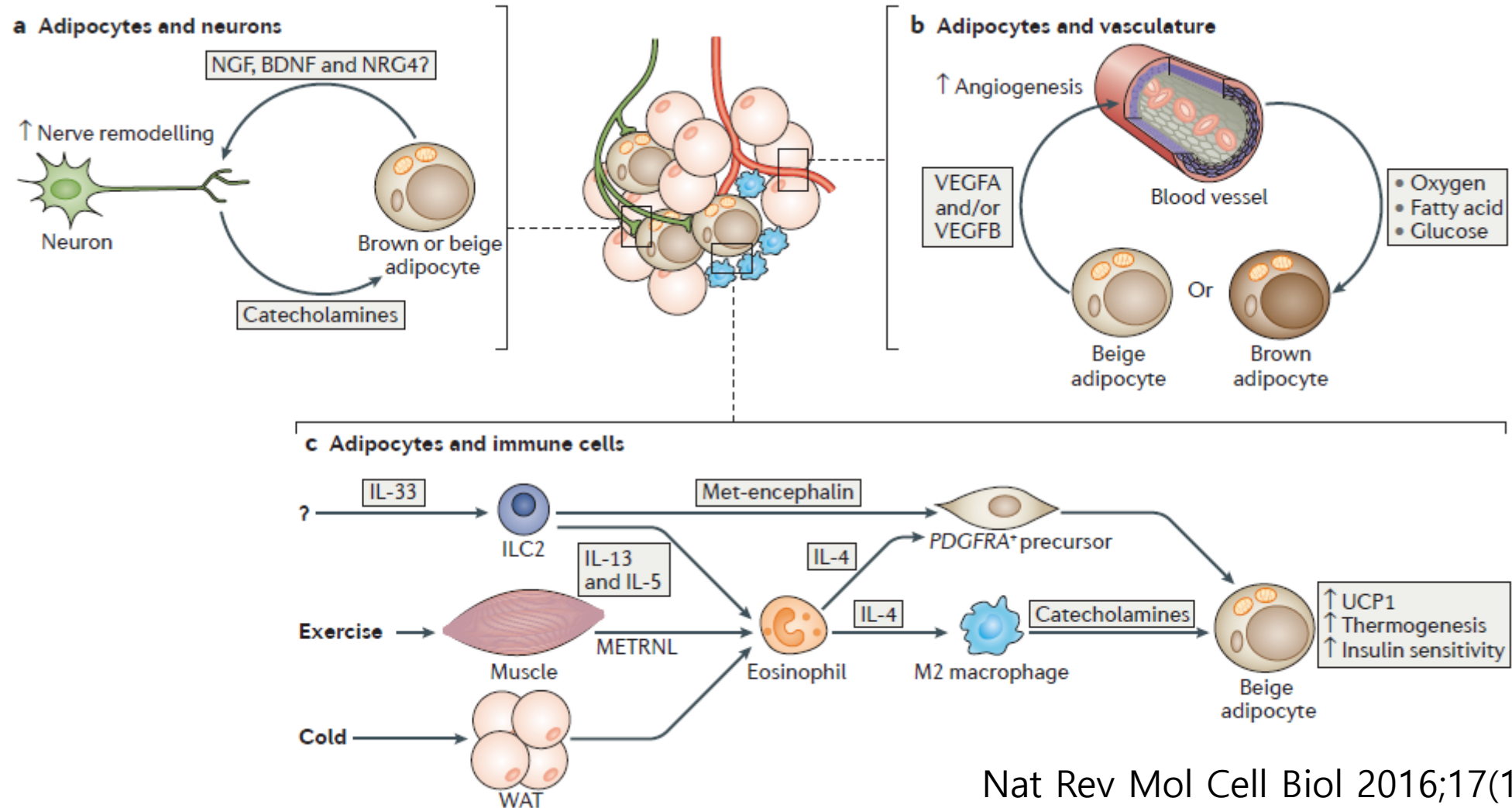
Associations of specific adipose tissue depots with metabolic consequences and distinct genetic loci







3 types of Adipocyte : brown, white and beige.



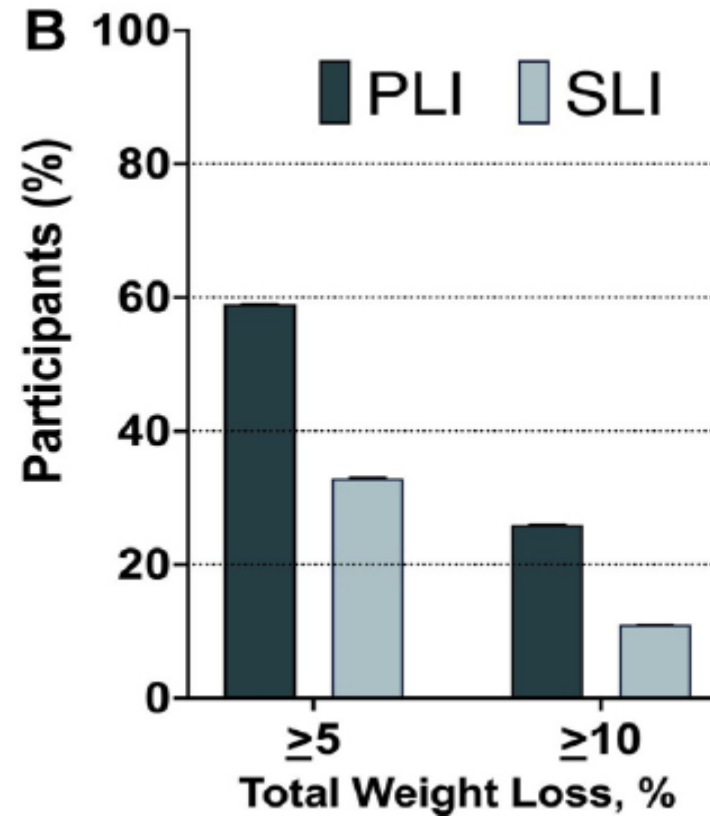
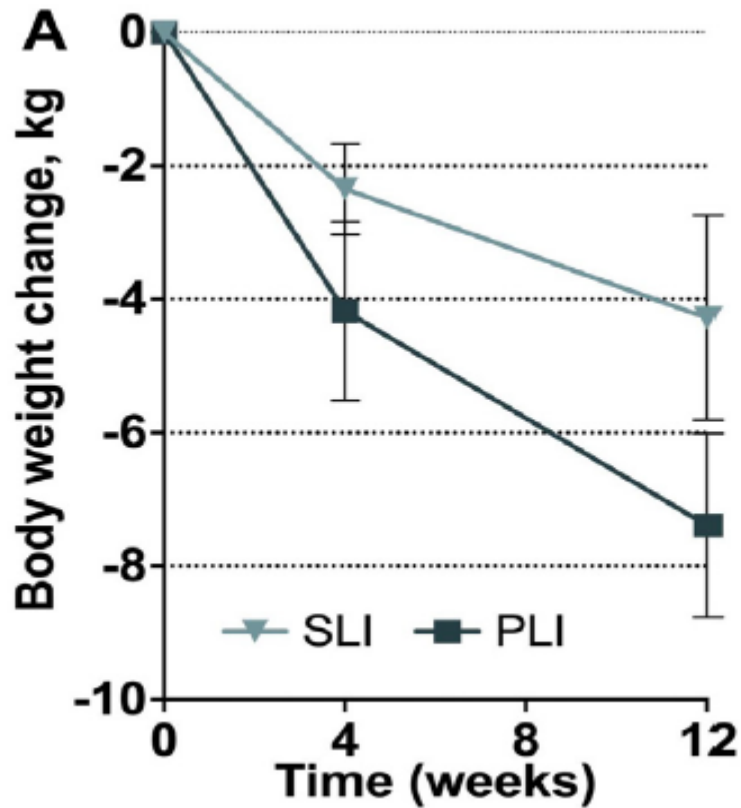
Crosstalk between brown and/or beige adipocytes and other adipose-resident cells



Functional Phenotypes of Obesity

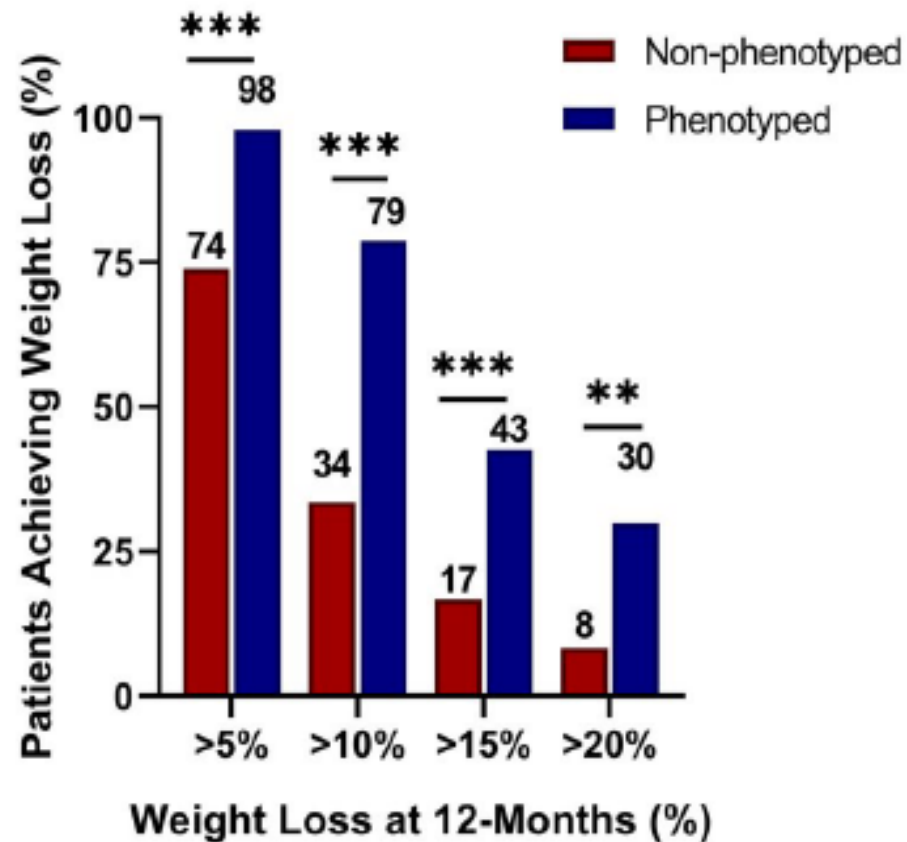
	Hungry brain 	Emotional hunger 	Hungry gut 	Slow burn 
Main obesity phenotype	Abnormal satiation*	Abnormal hedonic eating†	Abnormal satiety‡	Low predicted energy expenditure
Features (compared with a non-phenotype group)	<ul style="list-style-type: none"> • Consume 62% more calories before reaching fullness 	<ul style="list-style-type: none"> • 2-3 times higher anxiety levels 	<ul style="list-style-type: none"> • 31% faster gastric emptying rate 	<ul style="list-style-type: none"> • 12% lower predicted resting energy expenditure • Reduced muscle mass • Less active
Proposed targeted LIFESTYLE therapy	<ul style="list-style-type: none"> • Time-restricted eating 	<ul style="list-style-type: none"> • Low-calorie diet with intensive behavioural group therapy 	<ul style="list-style-type: none"> • Low-calorie diet with pre-meal protein supplements 	<ul style="list-style-type: none"> • Low-calorie diet with post-workout protein supplementation and high-intensity interval training
Proposed targeted PHARMACOTHERAPY	<ul style="list-style-type: none"> • Phentermine plus topiramate extended release 	<ul style="list-style-type: none"> • Oral naltrexone plus bupropion sustained release 	<ul style="list-style-type: none"> • Liraglutide 	<ul style="list-style-type: none"> • Phentermine

Total Body Weight loss (kg) in standard lifestyle intervention (SLI) and the phenotype-tailored lifestyle intervention (PLI).

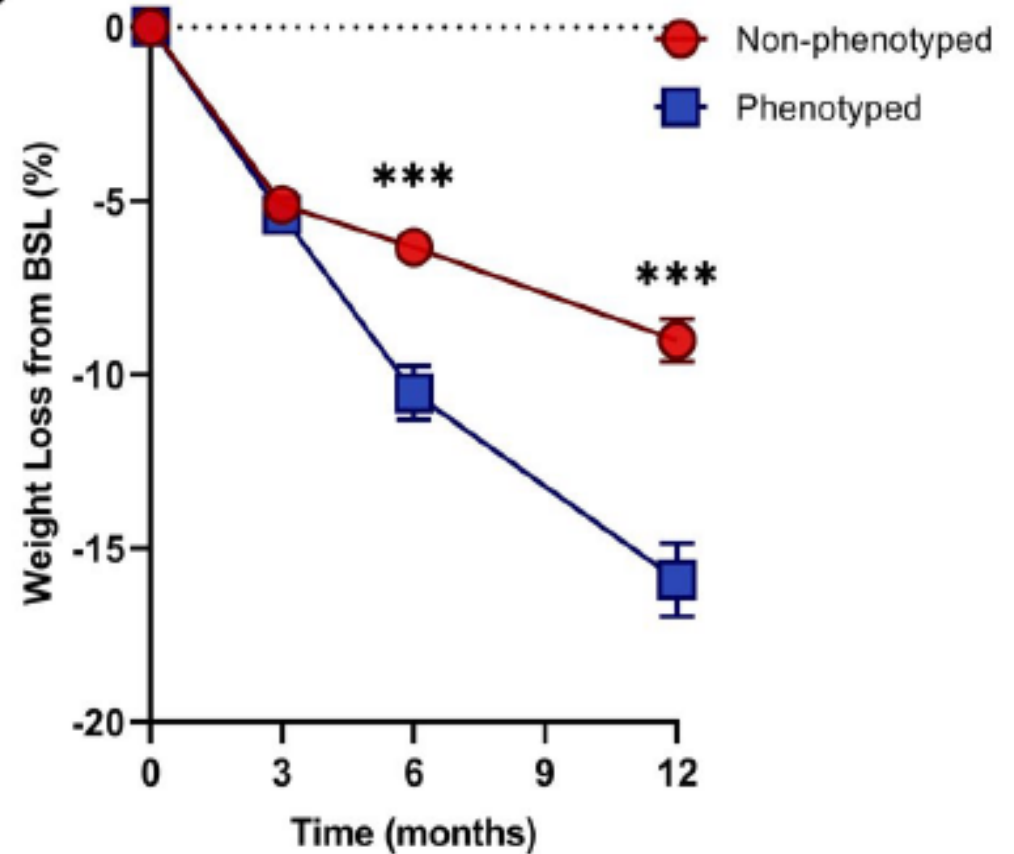


Phenotype Guided pharmacotherapy for obesity management improves weight loss outcomes

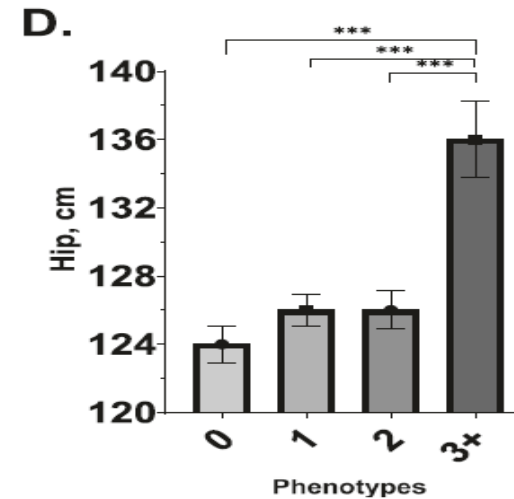
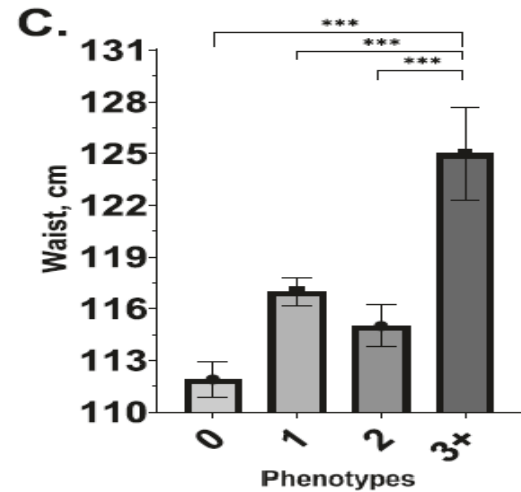
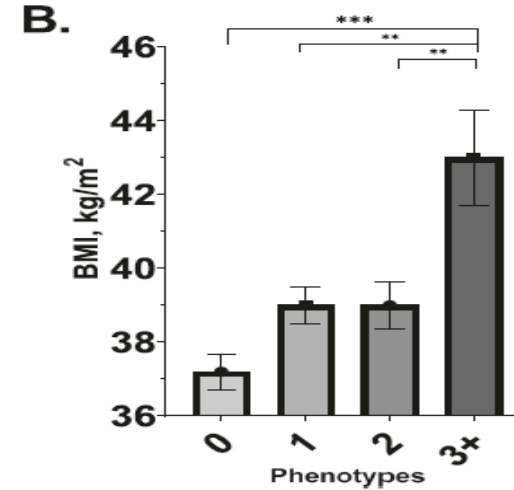
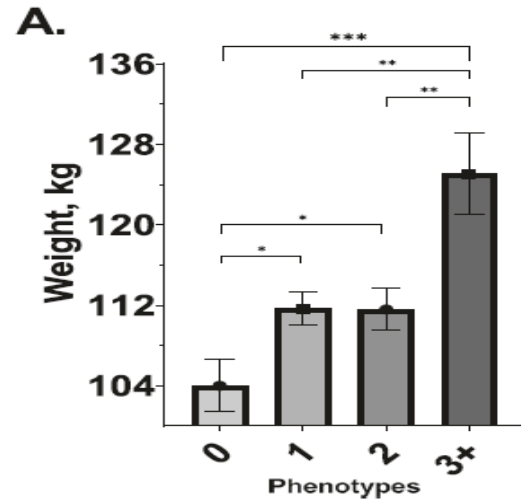
A



B



Cumulative effect of obesity phenotypes on body weight and body mass index



Clinical Obesity, Obesity addressed in clinical practice



Obesity Controversy

A Disease? vs **A Risk Factor?**



Obesity Controversy

A Disease? vs **A Risk Factor?**



Medical Treatment
Encouraging unhealthy behaviours
Financial Burden
Social responsibility
Social Stigma – disability



Life style intervention
Promoting healthy life
Financial freedom
Individual responsibility
Self-efficacy

The recognition of obesity as a disease

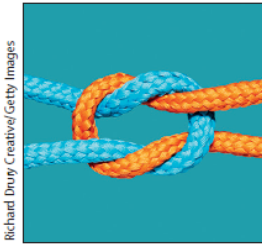
- associated with
 1. distinct pathophysiological alterations of tissues and organs
 2. discrete clinical signs and symptoms
 3. increased risk of secondary complication ,
 4. restrictions of daily activities.

NOT just with the number of BMI

CREATING A NEW HOLISTIC DIAGNOSTIC FRAMEWORK



Lancet Diabetes & Endocrinology Commission on the Definition and Diagnosis of Clinical Obesity



Published Online
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Obesity was first recognised as a disease by WHO in 1948, then between 2013 and 2022 by several medical societies and countries.¹⁻⁸ However, the notion that obesity is a disease and not merely a risk factor for other illnesses remains highly controversial, both within and beyond medical circles. This debate constitutes far more than arcane semantics, and seriously affects the provision of therapeutic strategies to improve health among people living with obesity.

On one side of the controversy, there is concern that defining obesity as a disease could have negative ramifications on individuals who have obesity and on society overall, by minimising the role of individual responsibility, thereby encouraging unhealthy behaviours and undermining efforts to tackle the increasing rates of obesity globally. Such arguments might, in part, reflect current intrinsic weight bias and stigma in our societies. Some experts also point to the fact that although a BMI in the traditional obesity range (ie, ≥ 27.5 kg/m² for people of Asian origin or >30.0 kg/m² for everyone else¹) is well known to be associated with increased risk of illness and death, a risk factor per se is not a disease. Indeed,

and overdiagnosis of obesity. In our opinion, the risk of overdiagnosis is a legitimate concern, especially for policy makers, because a blanket definition of obesity as a disease would classify approximately 30–40% of people in many nations as having this illness.⁹ This definition could render over a third of these populations suddenly eligible for claims of disability or expensive treatments. Such claims would effectively make obesity a financially and socially intractable issue. In summary, there is apprehension within and outside the medical profession that categorising obesity as a disease could unnecessarily medicalise the problem, undermining adherence to healthy lifestyles and potentially resulting in unwarranted use of drugs, medical technologies, and surgical procedures, with staggering economic and societal costs.

On the other side of the controversy, those who support the recognition of obesity as a disease cite evidence that the condition, like any other chronic disease state, is associated with distinct pathophysiological alterations of tissues and organs, discrete clinical signs and symptoms, increased risk

The 4 conditions of New diagnostic Framework for Clinical Obesity

- 1. New diagnostic tools should has a precise correlation with body fat and provide information about fat distribution**
- 2. New diagnostic tools should offer better prediction for obesity related consequences including mortality than BMI.**
- 3. New diagnostic tools should inform the genetic, metabolic, physiological or psychological contributions involved in the development of obesity.**
- 4. New diagnostic tools should be easy to apply assessment in clinical practice and medical research.**

57 years old Man

- Height 1.81 m
- Weight 87 Kg
- BMI 36.5 Kg/m²
- Waist Circ 100 cm
- Family History of
 - Acute MI, T2DM
 - Stroke
 - Dementia
 - Parkinsonism
 - Colon cancer
- Non-smoker
- Blood pressure 122/78 mmHg
- Glucose 89 mg/dl
- HbA1c 5.6%
- Lipid profile 206/146/69/108 mg/dl
- Mild fatty liver on abdominal us
- Past History of
 - Hepatitis B
 - Pulmonary Tuberculosis
 - Herpes Zoster
 - Exercise induced Asthma
 - Atopic dermatitis
- 10 units of alcohol once a week

Summary

- **The current diagnosis of obesity based on BMI has serious limitations for individuals in identifying excess adiposity and predicting obesity related consequences of them.**
- **Current clinical practice and health research is tied to BMI. There have been some shifts in looking at non-weight-related health indicators. However, we need more robust studies evaluating other health indicators beyond weight and BMI.**
- **The availability of these tools will help eliminate the need for BMI and promote individualized health assessment in future.**