

3° WORKSHOP CONGIUNTO SICOb – SID – SIO  
7 MARZO 2014

L'integrazione tra terapia medica e chirurgica nel  
trattamento del paziente obeso diabetico

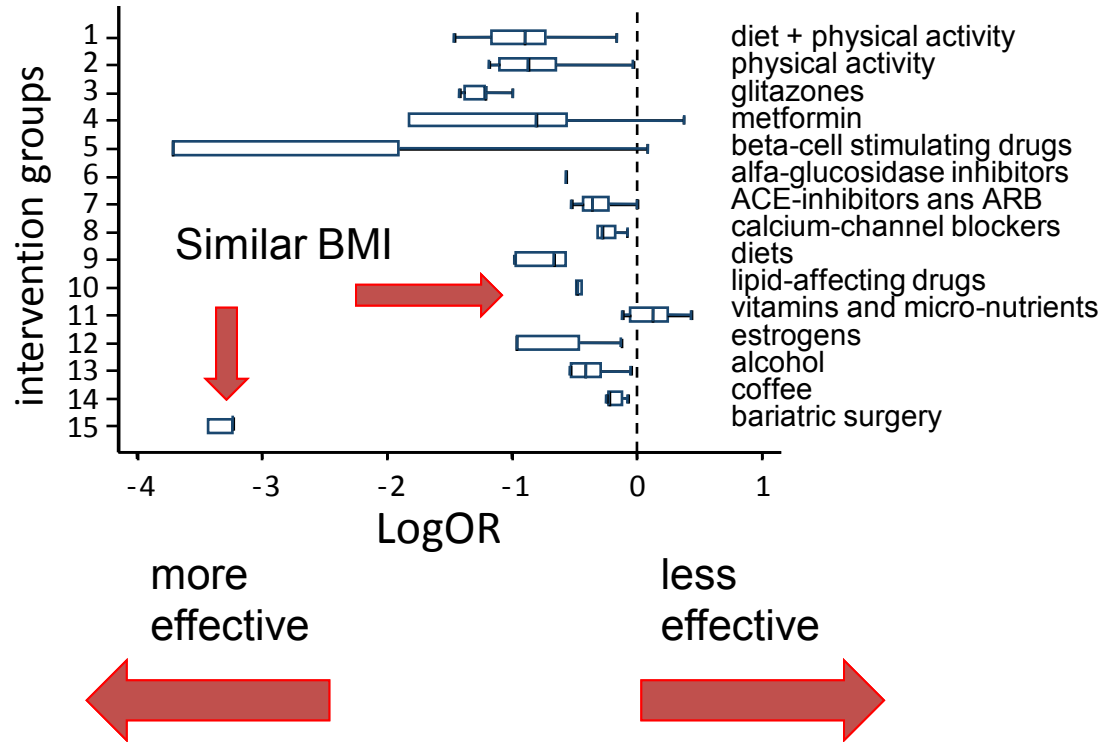
Recidive del diabete dopo terapia chirurgica  
La quantificazione del problema: quanto e dopo  
quanto



**antonio e. pontiroli**  
**clinica medica, DISS,**  
**università degli studi di milano,**  
**ospedale san paolo**



# prevention of type 2 diabetes; a systematic review and meta-analysis of different intervention strategies



*merlotti et al, diabetes obesity & metabolism (epub 2014)*

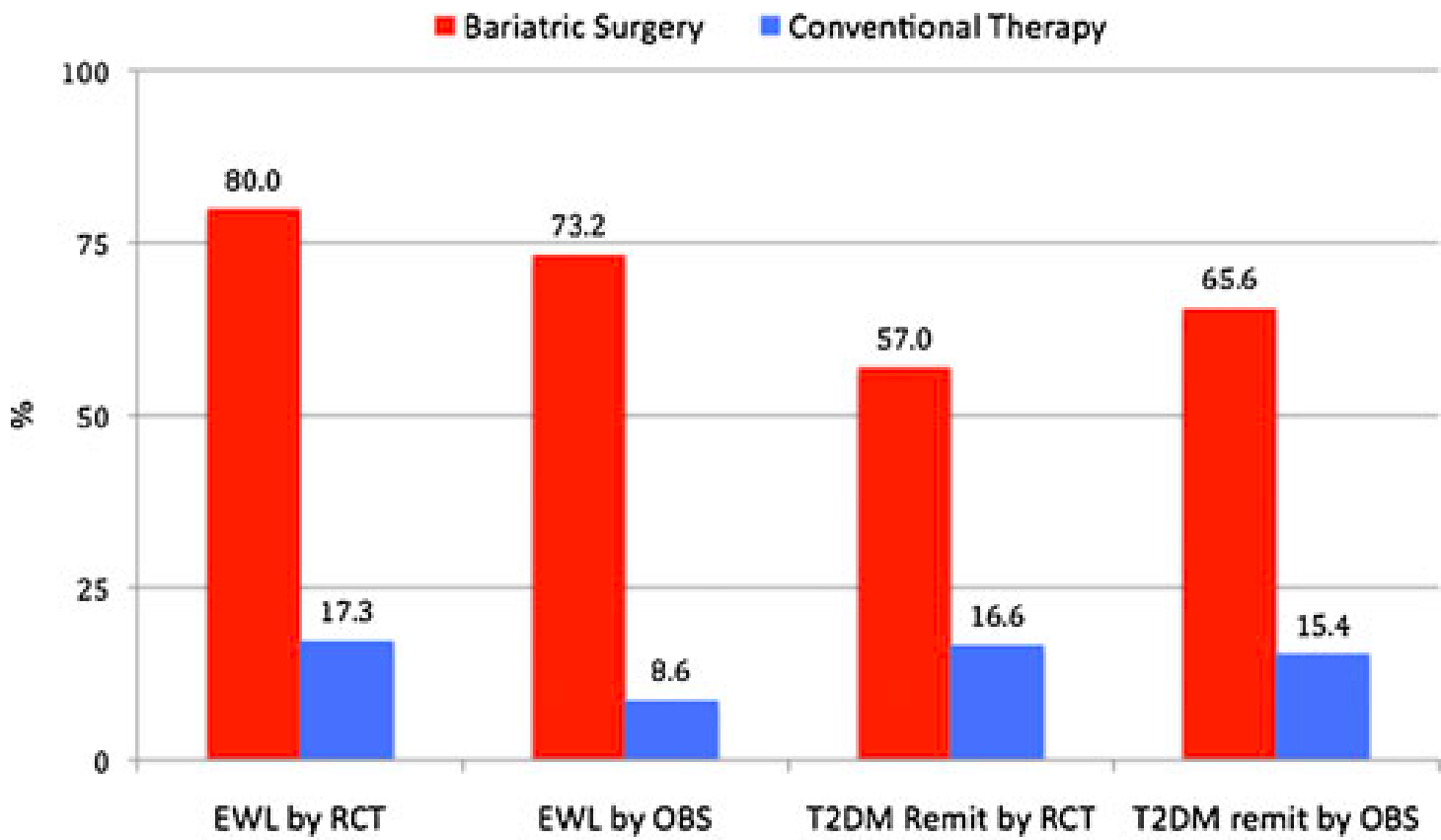
## bariatric surgery for obese type 2 diabetes: random studies

**Table 1** Analysis of the three randomized studies comparing surgery and medical treatment in resolution of type 2 diabetes mellitus.

	LAGB vs conventional treatment	RYGB or LSG vs intensive treatment	RYGB or BPD vs conventional treatment
Study reference	1	2	3
Numbers	30 + 30	50 + 50 + 50	20 + 20 + 20
BMI (kg/m <sup>2</sup> )	37.0	37.0	45.0
Criteria for T2DM resolution	FBG < 126 mg/dl HbA1c < 6.2%	HbA1c < 6.0%	FBG < 100 mg/dl HbA1c < 6.5%
Waist (cm)	114–116	114–116	125–126
Age (y)	47.0	49.8	43.5
Duration of T2DM (y)	<2	8	6
Weight loss	25.0% vs 1.5%	29% vs 25% vs 5.4%	33% vs 33% vs 5%
Resolution (without medications)	73% vs 13% nd	42% vs 37% vs 12% 29% vs 25% vs 0%	75% vs 95% vs 0% nd
Duration of study (y)	2	1	2
	<b>dixon (JAMA 2008)</b>	<b>schauer (NEJM 2012)</b>	<b>mingrone (NEJM 2012)</b>

LAGB = laparoscopic adjustable gastric banding; RYGB = gastric bypass; LSG = laparoscopic sleeve gastrectomy; BPD = biliopancreatic diversion; nd = not determined.

# Diabetes and Weight in Comparative Studies of Bariatric Surgery vs Conventional Medical Therapy: A Systematic Review and Meta-Analysis

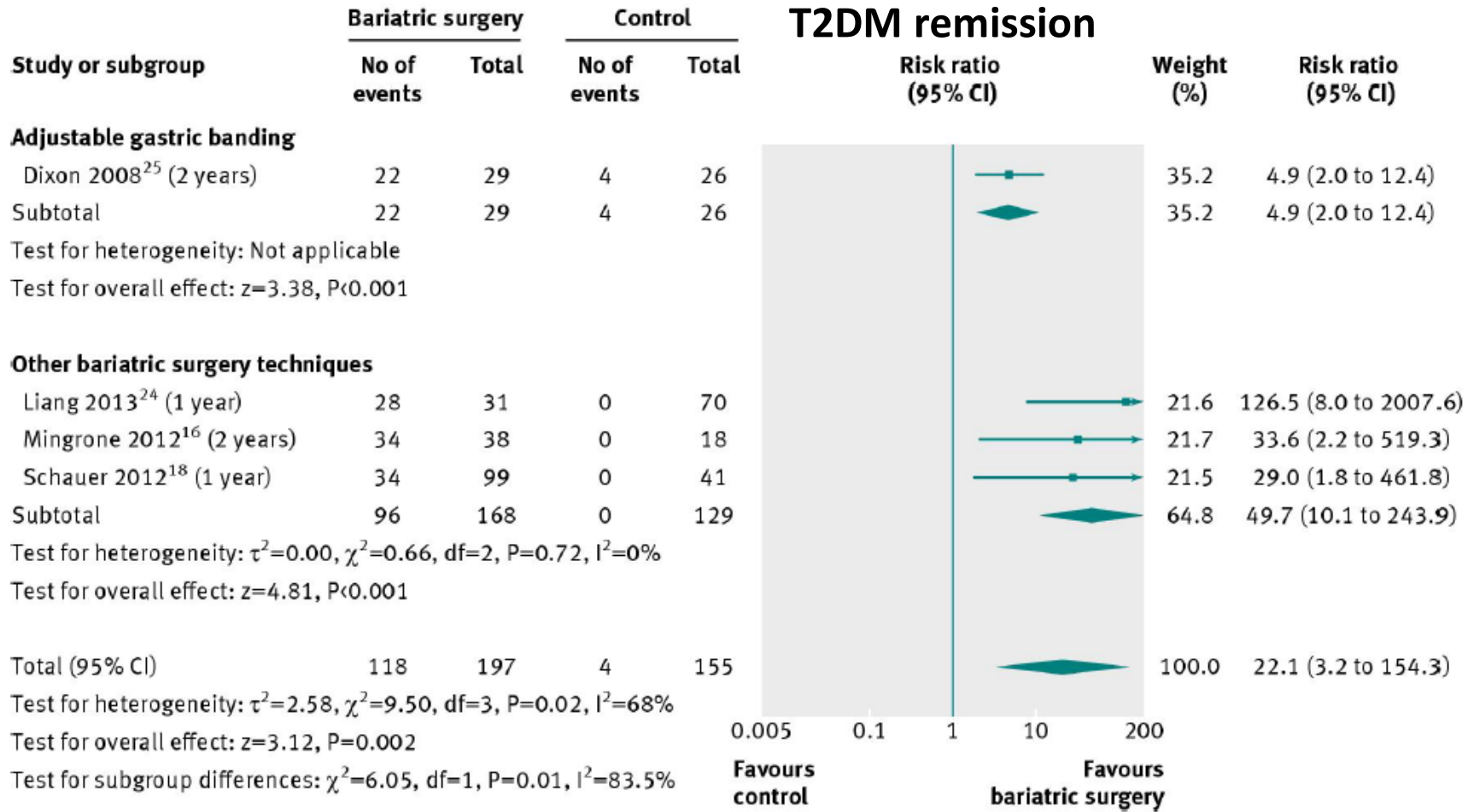


Conventional, conventional treatment; Surgery, combined bariatric surgical procedures; EWL, excess weight loss; RCT, randomized controlled trial; OBS, observational study.

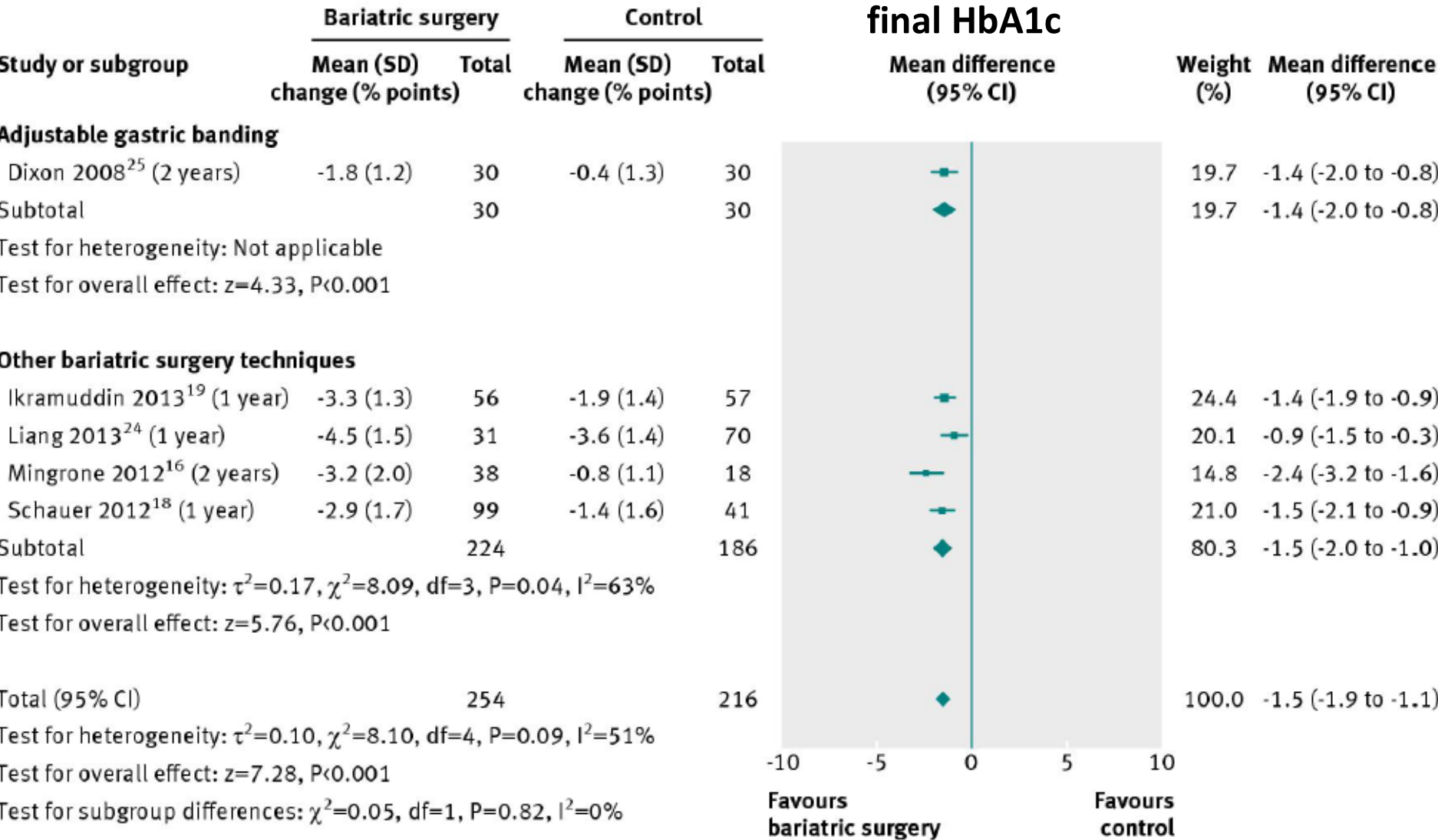
**total number 6,131; 3,076 bariatric surgery. 3,055 conventional treatment.**

*ribaric et al, obes surg 2014; 24: 437-455*

# Bariatric surgery versus non-surgical treatment for obesity: a systematic review and meta-analysis of randomised controlled trials

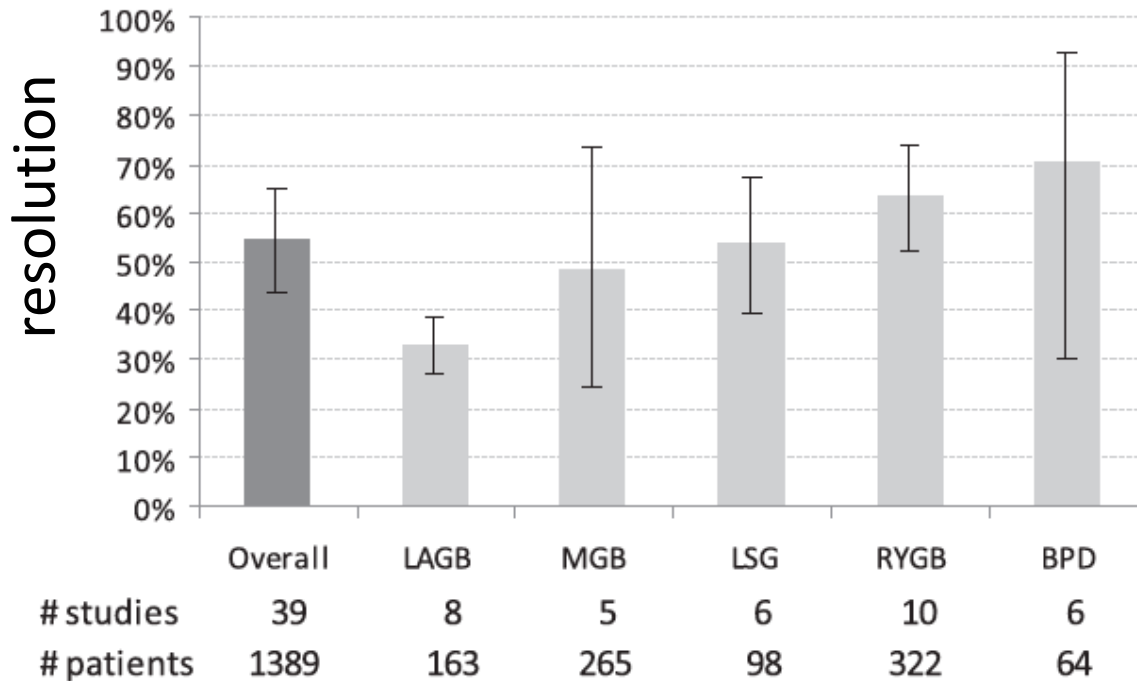


# Bariatric surgery versus non-surgical treatment for obesity: a systematic review and meta-analysis of randomised controlled trials



role of bariatric surgery as treatment for type 2 diabetes in patients who do not meet current NIH criteria: a systematic review and meta-analysis (BMI < 35 kg/m<sup>2</sup>).

*parikh et al, j am coll surg 2013; 217: 527-532*



3 RCTs (290 patients BMI 30-40, with or w/o T2DM) found surgery associated with greater weight loss (range, 14.4-24 kg) and glycemic control (range, 0.9-1.43 point in HbA1c) during 1 to 2 years of follow-up than nonsurgical treatment.

Observational studies (600 patients) and meta-analyses of nonsurgical therapies (> 300 RCTs) support this finding at 1 or 2 years.

*maggard-gibbons et al, JAMA 2013; 309: 2250-2261*

# Is diabetes cured by bariatric surgery?

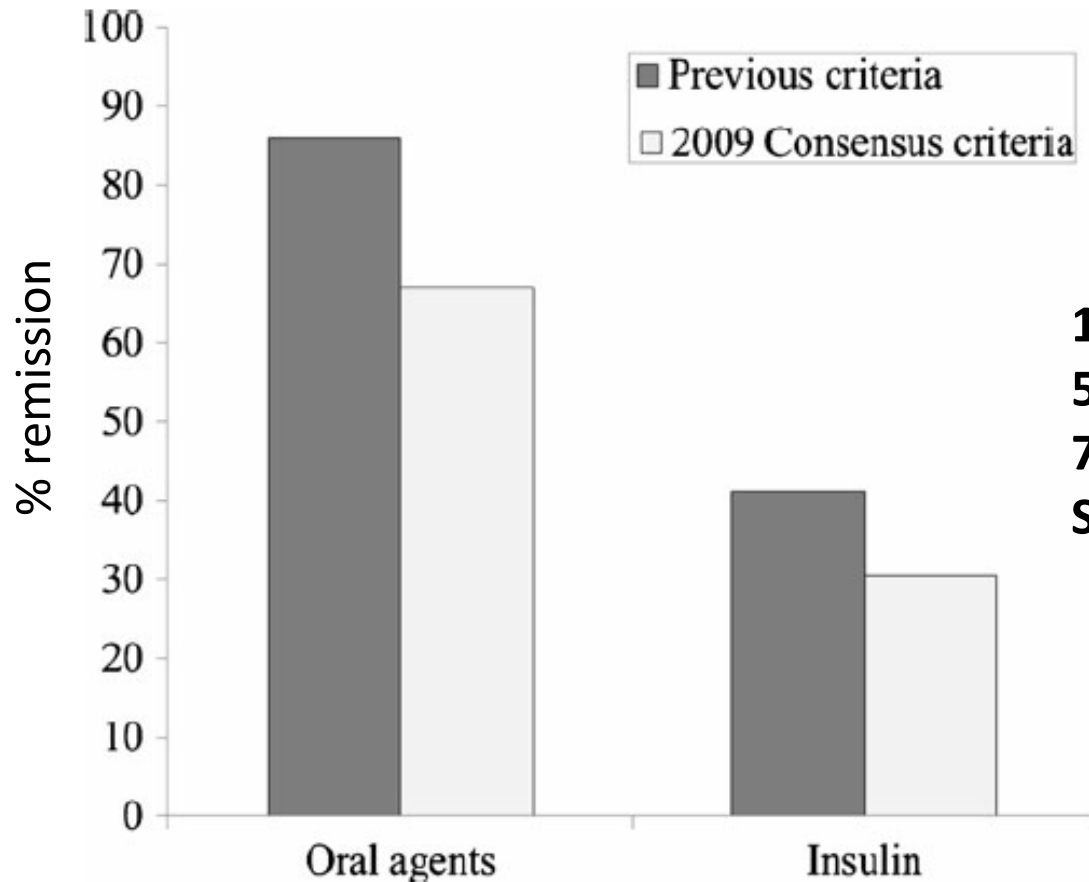
## Criteria for diabetes and diabetes resolution in bariatric surgery studies

1. Interview (1) and FBG < 126 mg/dl (2)
2. FBG < 126/100 mg/dl, HbA1c < 7/6%, no treatments (3-4)
3. OGTT (NGT, IFG/IGT, T2DM) (5)

*(1) Sjostrom, Obes Res 1999; 7: 477-484; (2) sjostrom, N Engl J Med 2004; 351: 2683-2693; (3) Buchwald, Amer J Med 2009; 122: 248-256; (4) Buse, Diabetes Care 2009; 32: 2133-2135; (5) Pontiroli, J Clin Endocrinol Metab 2002; 87: 3555-3561; Pontiroli, Diabetes Care 2005; 28: 2703-2709*



# Diagnosis of diabetes remission after bariatric surgery may be jeopardized by remission criteria and previous hypoglycemic treatment



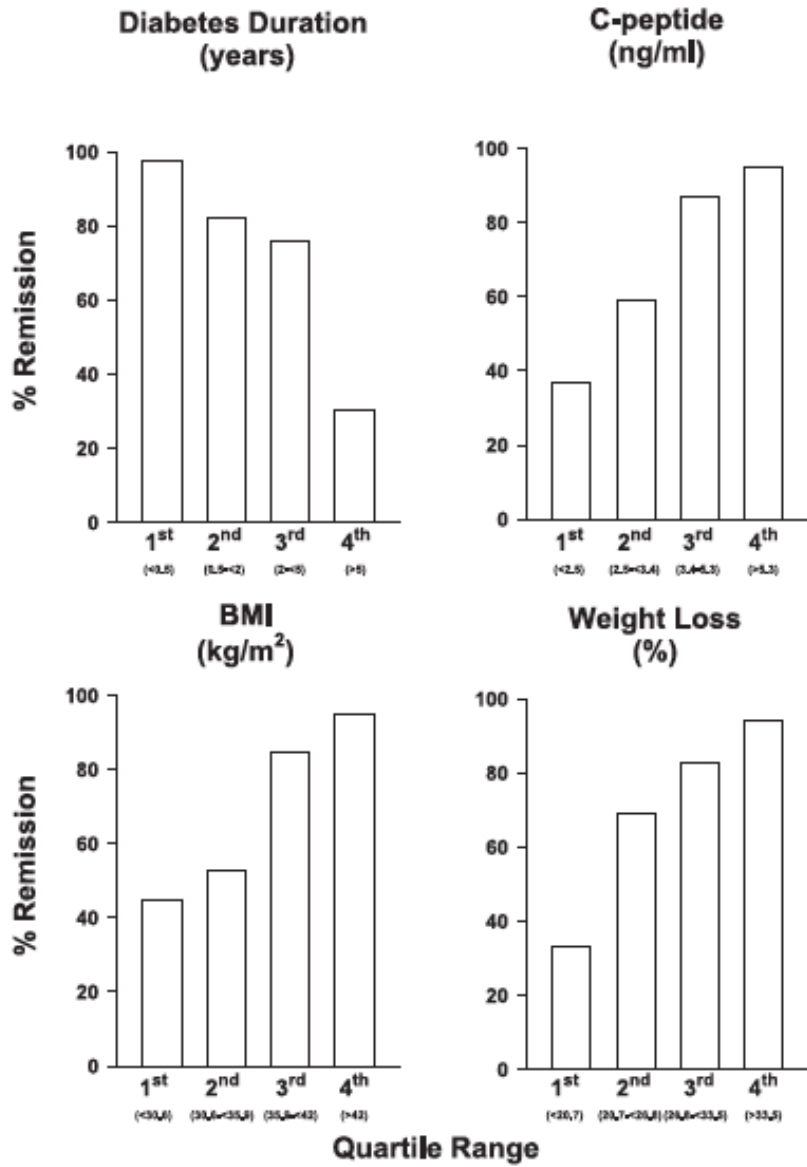
**141 T2DM patients:  
52 RYGB  
72 BPD  
SG 17**

Duration of diabetes, age, and female sex were associated to nonremission:  $10.3 \pm 9.4$  vs.  $4.7 \pm 3.8$  years,  $p < 0.001$ ;  $55.1 \pm 9.3$  vs.  $51.2 \pm 9.9$  years,  $p = 0.017$ ; 58.9 vs. 33.3%,  $p = 0.004$ , respectively

***ramos-levi et al. obes surg 2013; 23: 1520-1526***

# Predicting the glycemic response to gastric bypass surgery in patients with type 2 diabetes

154 patients,  
104 on remission  
at 1 year



# Factors associated with T2DM remission - recurrence

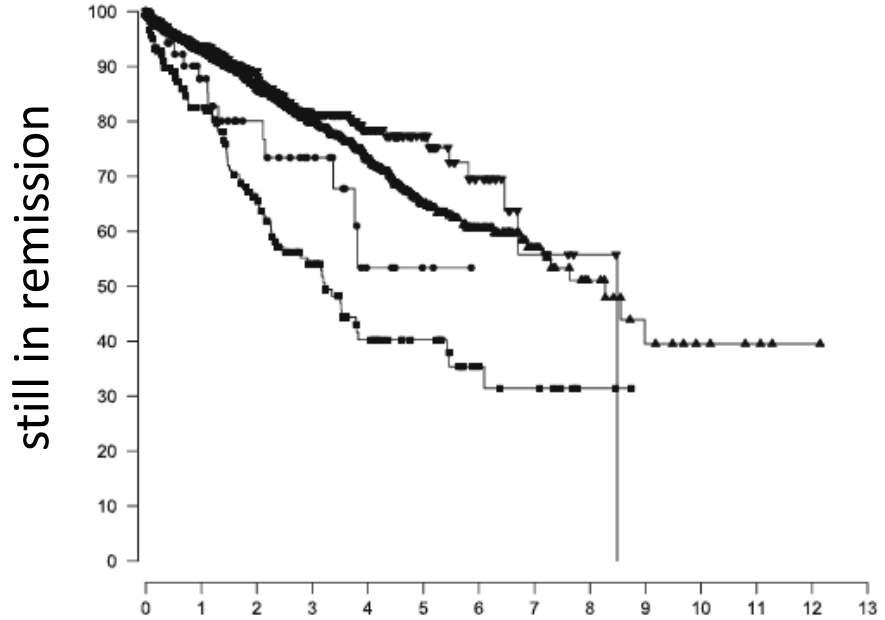
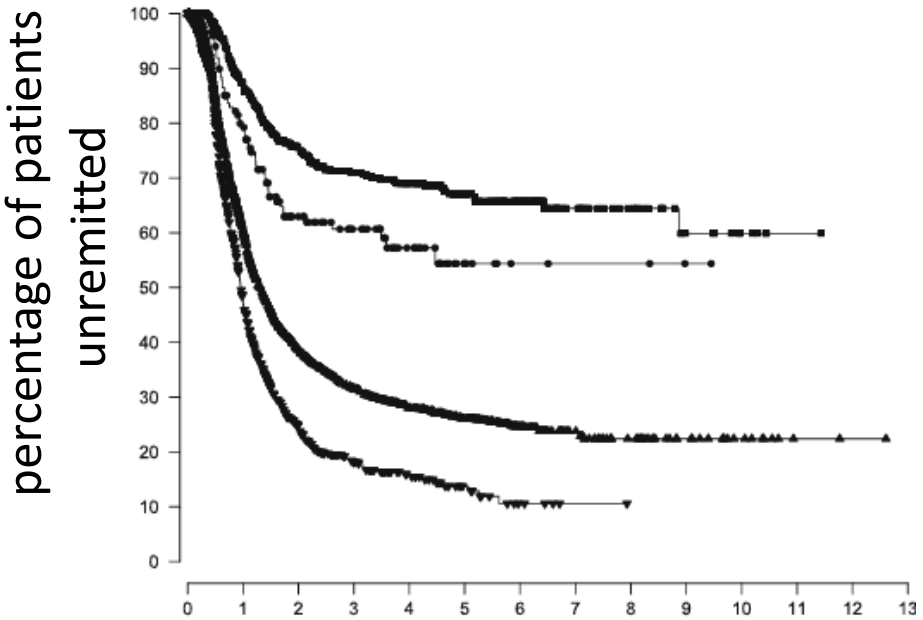
subjects	surgery	remission	recurrence
42 (1)	RYGB	64% improved	24% recurred - Low BMI- poor weight loss – high FBG
177 (2)	RYGB		43% -previous treatments
97 (3)	RYGB	Older, high BMI, use of drugs	
46 (4)	LAGB + RYGB	Duration < 4y, HbA1c < 7.1, BMI < 50	
126 (5)	RYGB	C peptide	
153 (6)	RYGB + LSG		Older age, insulin use, weight regain
269 (7)	RYGB		Weight regain

(1) digiorgi et al SOARD 2010; 6. 249-253; (2) chikunguwo et al SOARD 2010; 6: 254-259; (3) yamaguchi et al, surg endosc 2012; 26: 2843-2847;(4) robert et al obes surg 2013; 23: 770-775; (5) aarts et al obes surg 2013; 23: 867-873; (6) jmenez et al ann surg 2012; 256: 1023-1029; (7) campos et al ABCD 2013; 26 (suppl 1): 57-62

# a multisite study of long-term remission and relapse of T2DM following gastric bypass

4,434 patients with T2DM,  
2,254 with clinical remission

- insulin, HbA1c > 6.5%
- Insulin, HbA1c < 6.5%
- ▲ no-insulin, HbA1c > 6.5%
- ▼ no-insulin, HbA1c < 6.5%



years since surgery

**predictors of relapse: high HbA1c, oral agents or insulin,  
low weight loss, duration of T2DM > 5 years**

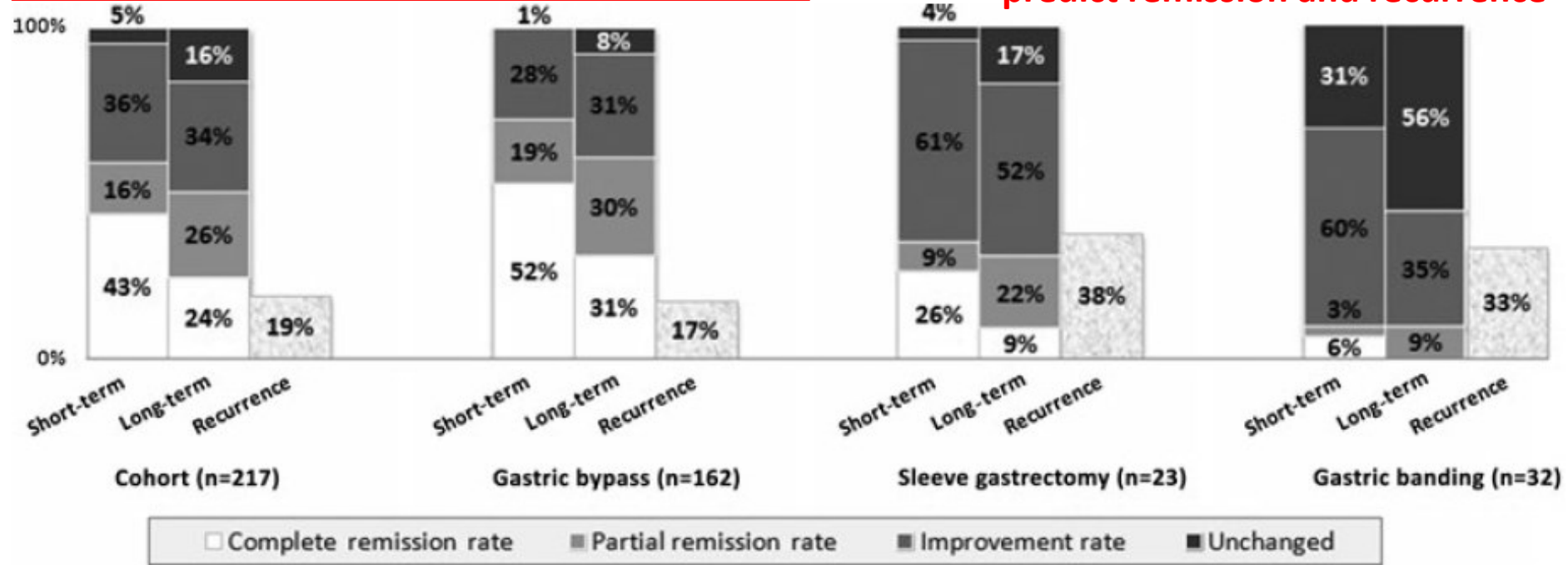
# Can diabetes be surgically cured? Long-term metabolic effects of bariatric surgery in obese patients with type 2 diabetes mellitus

	Whole Cohort	Gastric Bypass	Sleeve Gastrectomy	P1	Gastric Banding	P2
Total weight loss (%)						
Short-term	27.6 ± 10.1	30.9 ± 8.3	21.2 ± 10.6	<0.001	16.5 ± 7.6	0.068
Long-term	25.4 ± 11.9	28.1 ± 10.9	22.2 ± 9.3	0.015	13.2 ± 10.7	0.002
EWL (%)						
Short-term	60.3 ± 24.3	66.8 ± 20.4	49.7 ± 32.5	0.029	37.0 ± 17.8	0.112
Long-term	54.9 ± 26.7	60.5 ± 24.6	49.5 ± 24.9	0.047	29.5 ± 23.4	0.004

All values are mean ± SD.

Short-term: 1-2 yrs after surgery; Long-term: 5 yrs or more after surgery.  
 P1: gastric bypass vs sleeve gastrectomy; P2: sleeve gastrectomy vs gastric banding.

**T2DM duration and weight loss predict remission and recurrence**



**long-term control rates of HDL, LDL, TG, AH were 73%, 72%, 80%, and 62%, respectively.  
 diabetic nephropathy regressed (53%) or stabilized (47%).**

The future:

Proper identification of candidates

**Avoid dichotomy between surgery and drugs**

(Expectations of patients and clinicians may have to be adjusted as regards remission of type 2 diabetes after bariatric surgery, Pournaras, Br J Surg 2012; 99: 100-103)

**The effect of bariatric surgery can be of limited time value, but other effects can last longer**