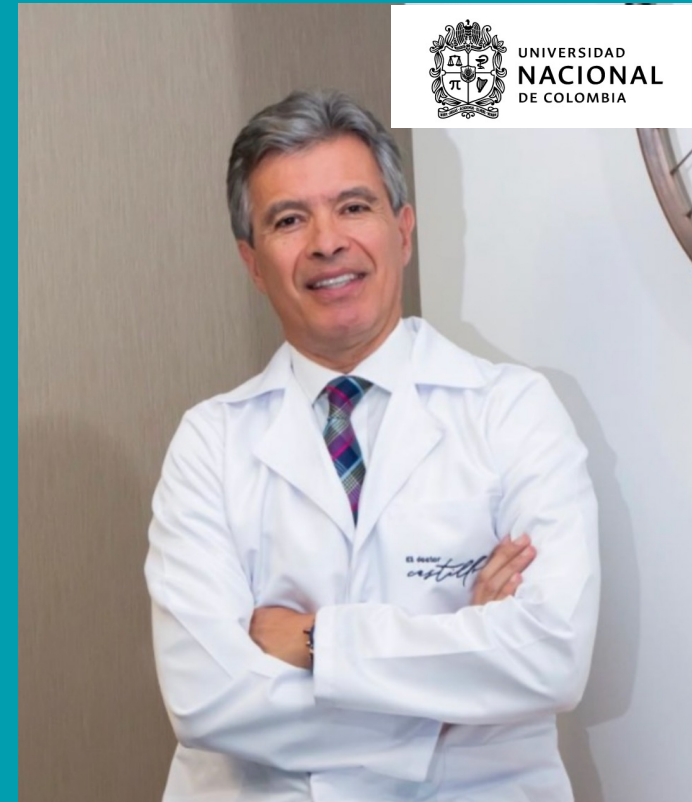


Mas allá de la glucosa: el papel de la insulina en la salud muscular del paciente viviendo con diabetes

# Castillo Jorge

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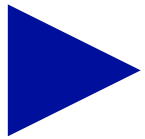
Especialista en Endocrinología  
Los Cobos Medical Center  
Bogotá, Colombia



# Conflicto de interés

- ▶  Esta es una conferencia patrocinada por laboratorios **Sanofi**
- ▶  Su contenido es producto de información científica no influenciada por el patrocinador
- ▶  He recibido honorarios como speaker de Amgen, Astra Zeneca, BD, Boeringher, Diabetrics, Euroetika, Gilead, Merck Serono, Merck Sharp and Dhome, Lilly, Novo Nordisk, Pharmatech, Pfizer, PTC, Procaps, Roche, Sanofi, Servier, Tecnofarma.

# Conferencia disponible en...



[www.eldoctorcastillo.com](http://www.eldoctorcastillo.com)





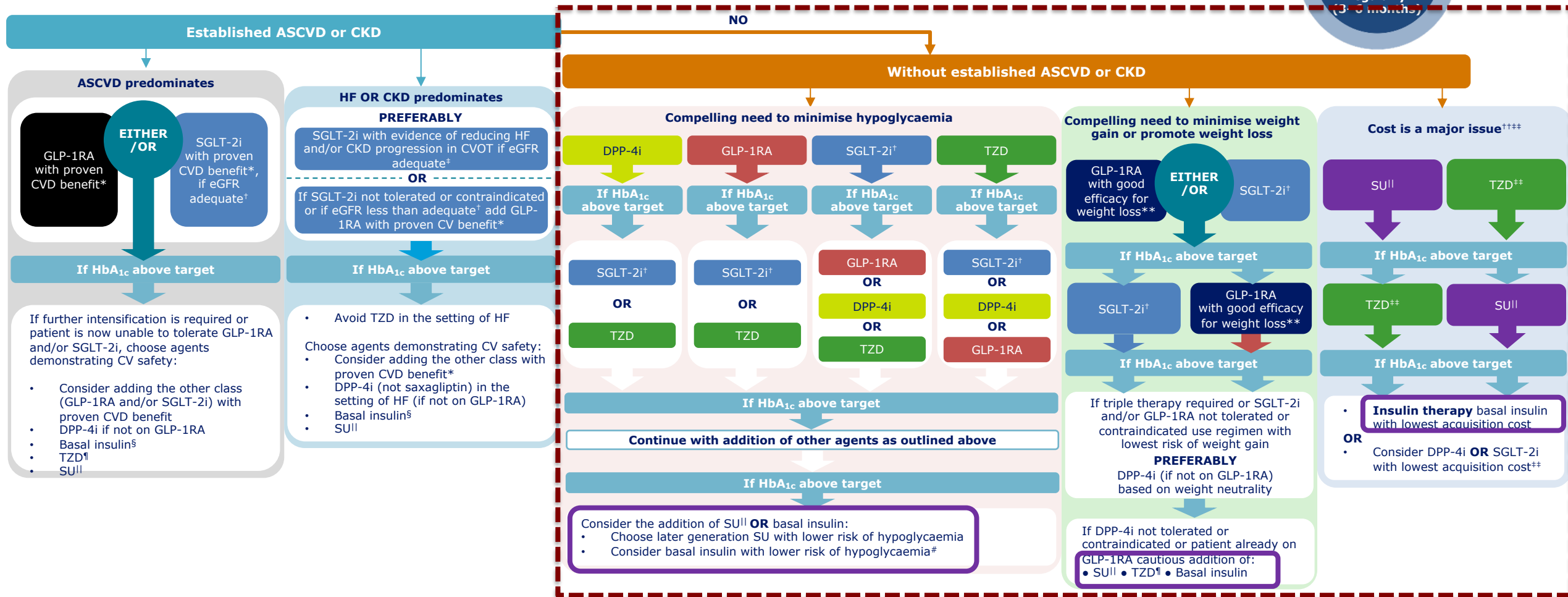
• **Lo que deberíamos saber de la Insulina**



@eldoctorcastillo

**FIRST-LINE THERAPY IS METFORMIN AND COMPREHENSIVE LIFESTYLE (INCLUDING WEIGHT MANAGEMENT AND PHYSICAL ACTIVITY)  
IF HbA<sub>1c</sub> ABOVE TARGET PROCEED AS BELOW**

To avoid clinical inertia reassess and modify treatment regularly (3-6 months)

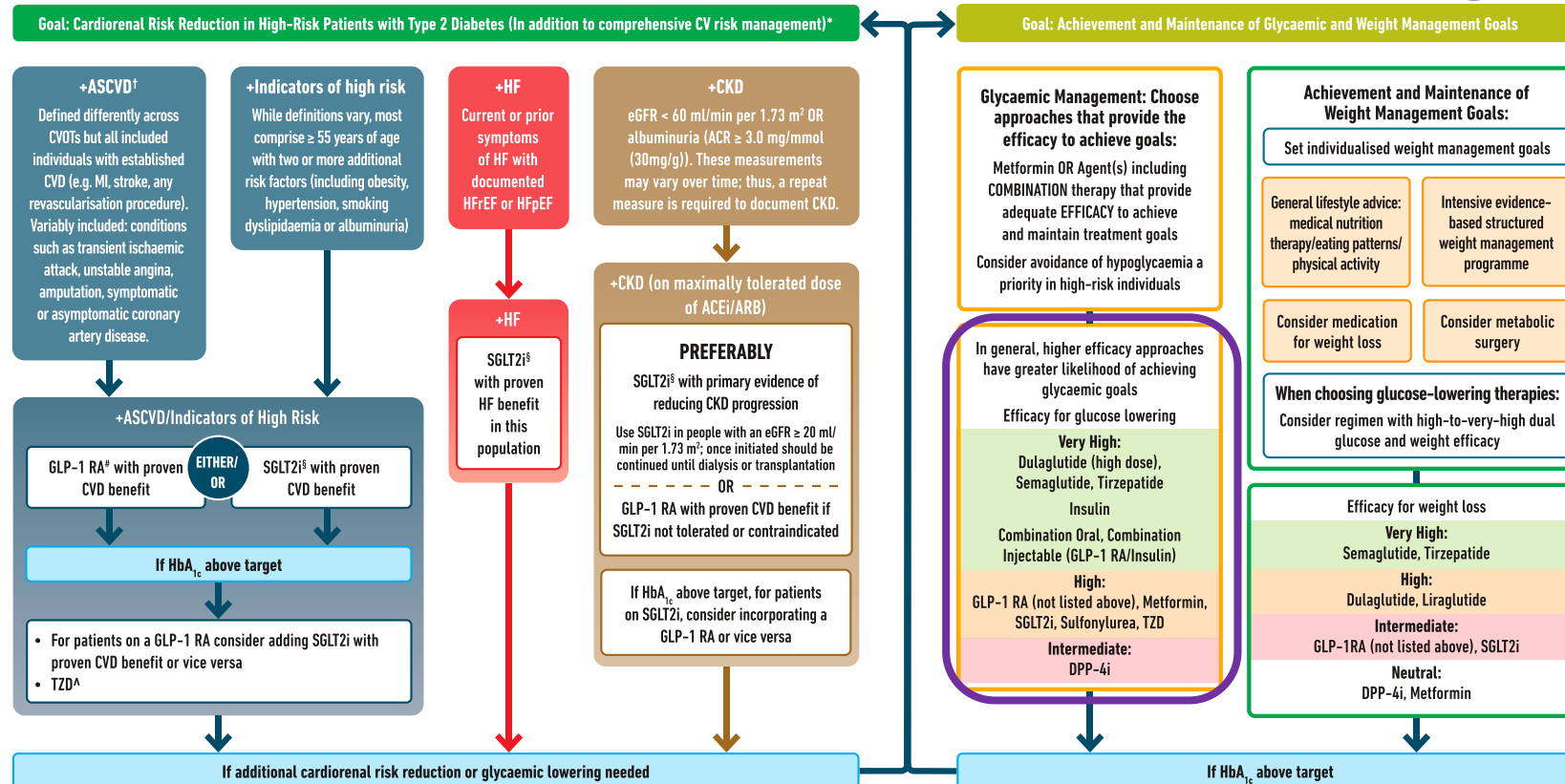


\*Proven CVD benefit means it has label indication of reducing CVD events. For GLP-1RA strongest evidence for liraglutide>semaglutide>exenatide extended release. For SGLT-2i evidence modestly stronger for empagliflozin>canagliflozin;

\*\*\*Semaglutide>liraglutide>dulaglutide>exenatide>lixisenatide; \*\*If no specific comorbidities (i.e. no established CVD, low risk of hypoglycaemia and lower priority to avoid weight gain or no weight-related comorbidities); ++Consider country- and region-specific cost of drugs. In some countries, TZDs relatively more expensive and DPP-4i relatively cheaper

## USE OF GLUCOSE-LOWERING MEDICATIONS IN THE MANAGEMENT OF TYPE 2 DIABETES

HEALTHY LIFESTYLE BEHAVIOURS; DIABETES SELF-MANAGEMENT EDUCATION AND SUPPORT (DSMES); SOCIAL DETERMINANTS OF HEALTH (SDOH)



\* In people with HF, CKD, established CVD or multiple risk factors for CVD, the decision to use a GLP-1 RA or SGLT2i with proven benefit should be independent of background use of metformin; † A strong recommendation is warranted for people with CVD and a weaker recommendation for those with indicators of high CV risk. Moreover, a higher absolute risk reduction and thus lower numbers needed to treat are seen at higher levels of baseline risk and should be factored into the shared decision-making process. See text for details; ^ Low-dose TZD may be better tolerated and similarly effective; § For SGLT2i, CV/renal outcomes trials demonstrate their efficacy in reducing the risk of composite MACE, CV death, all-cause mortality, MI, HFrEF and renal outcomes in individuals with T2D with established/high risk of CVD; # For GLP-1 RA, CVOTs demonstrate their efficacy in reducing composite MACE, CV death, all-cause mortality, MI, stroke and renal endpoints in individuals with T2D with established/high risk of CVD.

Identify barriers to goals:

- Consider DSMES referral to support self-efficacy in achievement of goals
- Consider technology (e.g. diagnostic CGM) to identify therapeutic gaps and tailor therapy
- Identify and address SDOH that impact on achievement of goals



In general, higher efficacy approaches have greater likelihood of achieving glycaemic goals

Efficacy for glucose lowering

**Very High:**

Dulaglutide (high dose),  
Semaglutide, Tirzepatide

Insulin

Combination Oral, Combination  
Injectable (GLP-1 RA/Insulin)

**High:**

GLP-1 RA (not listed above), Metformin,  
SGLT2i, Sulfonylurea, TZD

**Intermediate:**

DPP-4i

## Glycemia Reduction in Type 2 Diabetes — Glycemic Outcomes

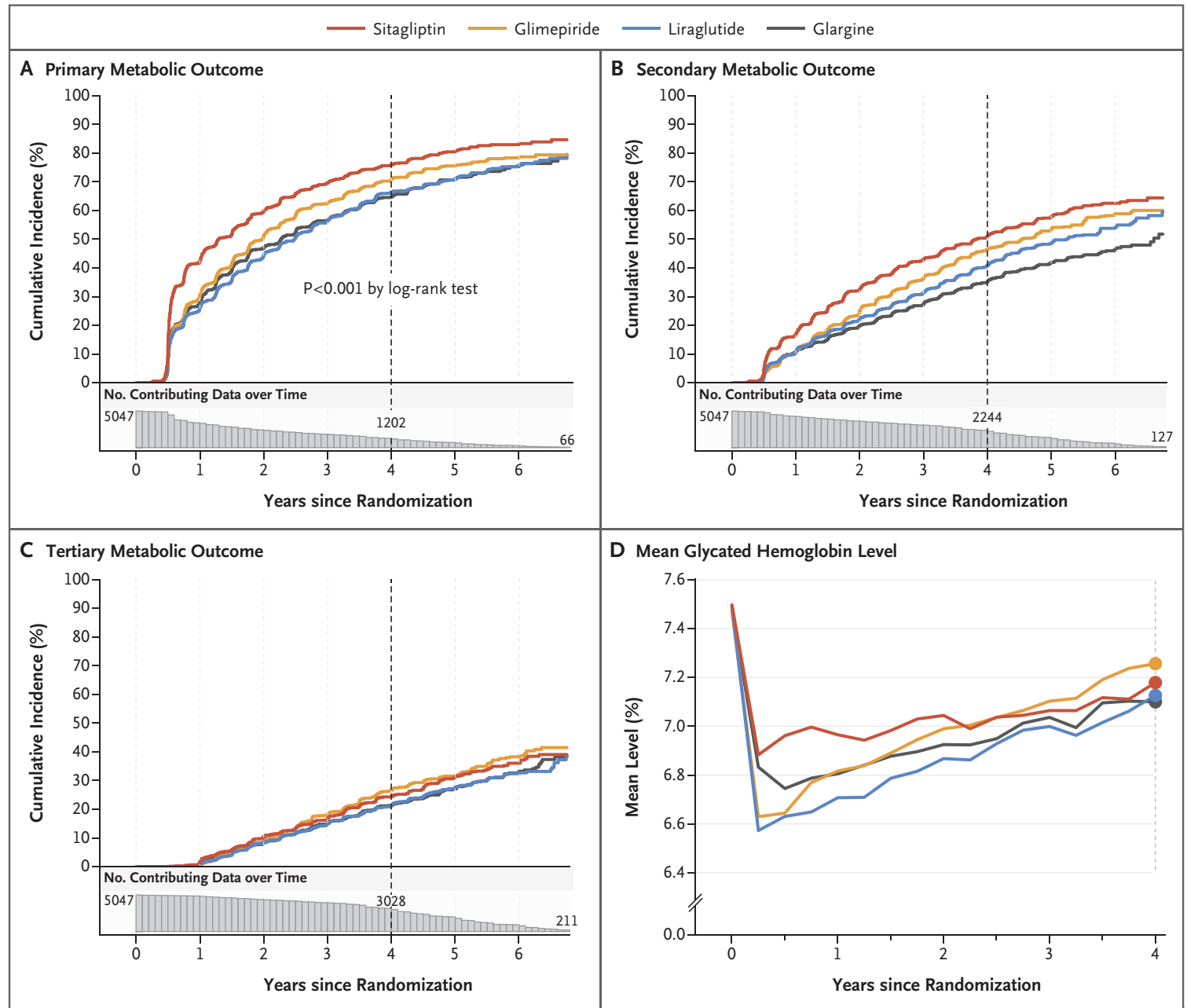
The GRADE Study Research Group\*

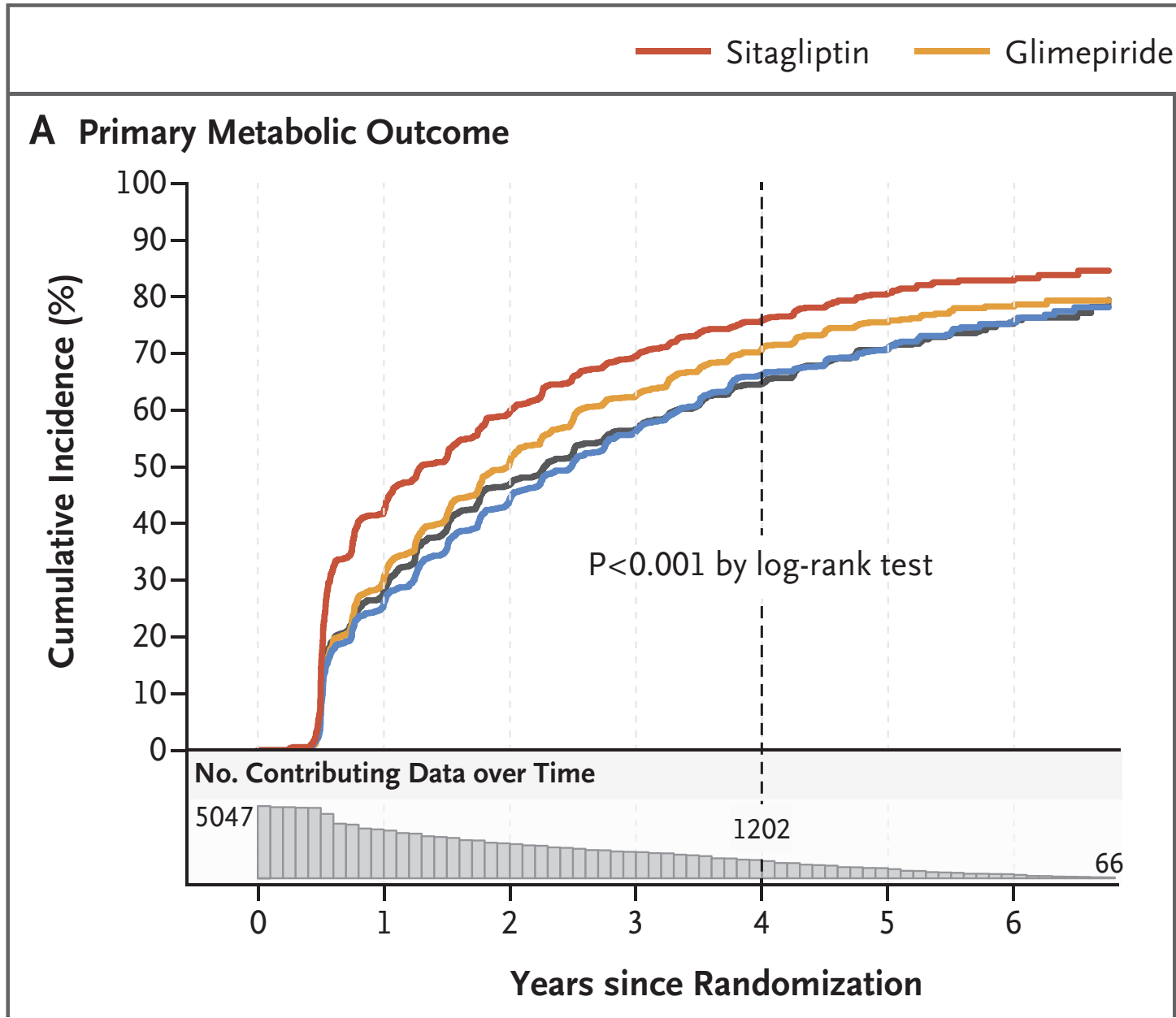
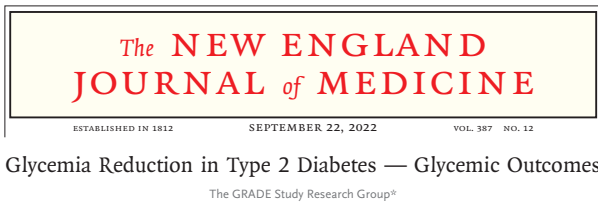
**Table 2. Primary and Secondary Metabolic Outcomes.\***

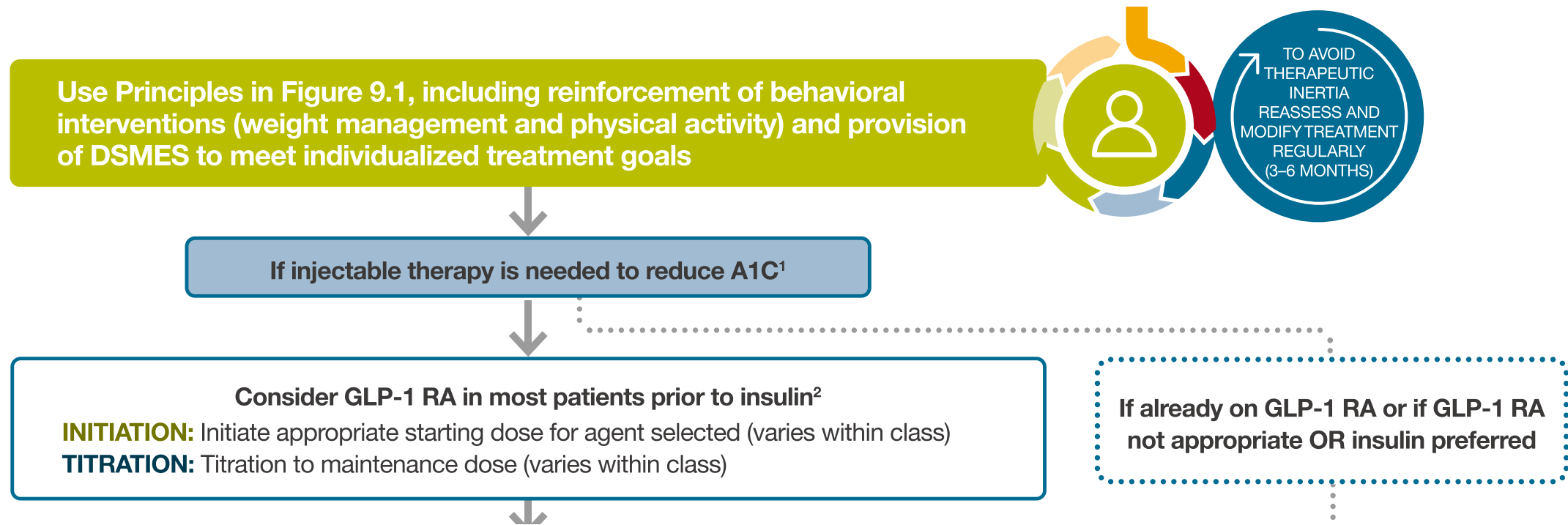
Outcome	Glargine (N = 1263)	Glimepiride (N = 1254)	Liraglutide (N = 1262)	Sitagliptin (N = 1268)
<b>Primary metabolic outcome†</b>				
Participants — no. (%)	852 (67.4)	908 (72.4)	860 (68.2)	981 (77.4)
Rate/100 participant-yr (95% CI)	26.5 (24.8–28.4)	30.4 (28.4–32.4)	26.1 (24.4–27.9)	38.1 (35.8–40.6)
Pairwise hazard ratios (95% CI)				
Glargine	—	0.89 (0.81–0.98)‡	1.02 (0.93–1.12)	0.71 (0.64–0.78)§
Glimepiride	—	—	1.15 (1.04–1.27)¶	0.79 (0.72–0.88)§
Liraglutide	—	—	—	0.69 (0.63–0.76)§
Hazard ratio in the treatment group as compared with all other treatments combined (95% CI)	0.87 (0.80–0.94)§	1.01 (0.93–1.09)	0.84 (0.78–0.91)§	1.37 (1.27–1.48)§



Glycemia Reduction in Type 2 Diabetes — Glycemic Outcomes  
The GRADE Study Research Group\*







# Cúando Insulinizar

**1**

**HbA1c**

10%  
> 300 mg/dl

**2**

**HbA1c**

9%  
Síntomas  
clásicos

**5**

**ADOs**

2 o 3 sin  
llegar a  
metas

**3**

**Estados  
Catabólicos**

Estado Hiperosmolar  
Cetoacidosis  
Urgencias

**4**

**Condiciones  
especiales**

Embarazo  
Hospitalización

# HIPOGLUCEMIA Y PESO

en diabetes tipo 1





ORIGINAL PAPER

 THE INTERNATIONAL JOURNAL OF  
**CLINICAL PRACTICE**

## **How much is too much? Outcomes in patients using high-dose insulin glargine**

T. Reid,<sup>1</sup> L. Gao,<sup>2</sup> J. Gill,<sup>3</sup> A. Stuhr,<sup>3</sup> L. Traylor,<sup>3</sup> A. Vlajnic,<sup>3</sup> A. Rhinehart<sup>4</sup>



# How much is too much? Outcomes in patients using high-dose insulin glargine

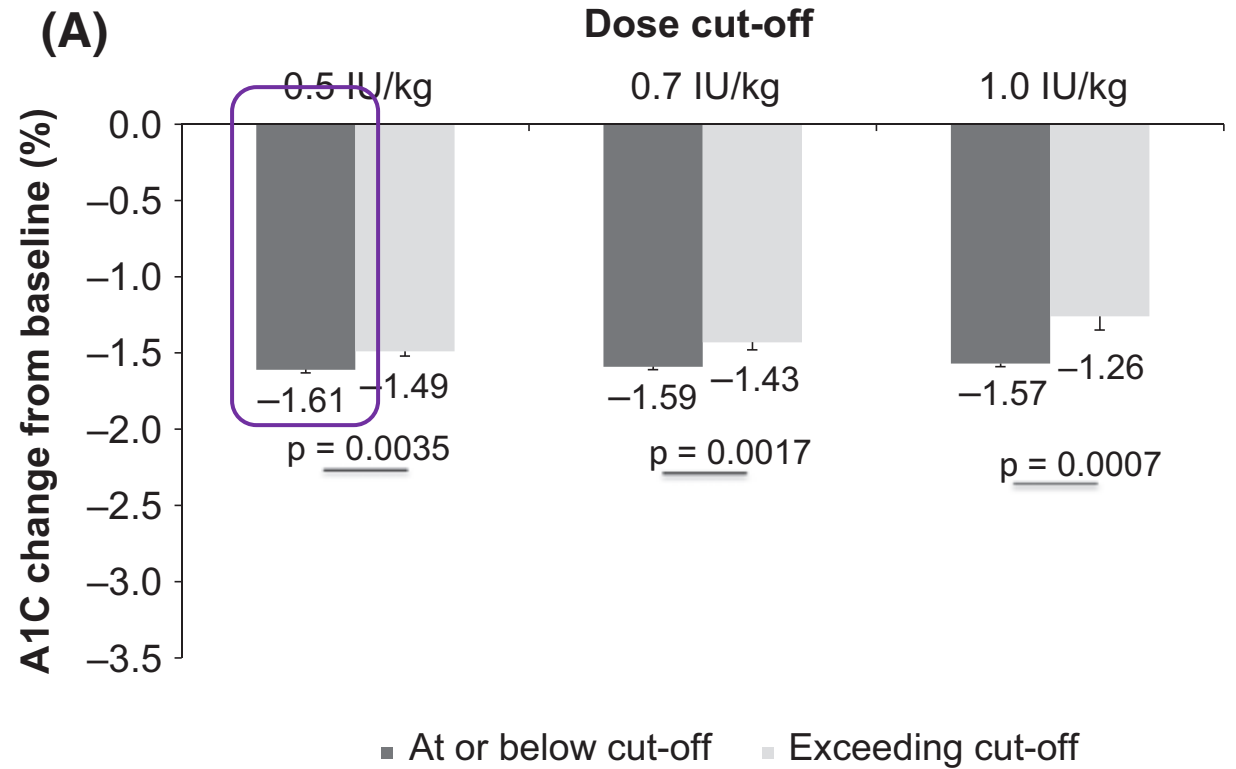
T. Reid,<sup>1</sup> L. Gao,<sup>2</sup> J. Gill,<sup>3</sup> A. Stuhr,<sup>3</sup> L. Traylor,<sup>3</sup> A. Vlainic,<sup>3</sup> A. Rhinehart<sup>4</sup>

0.5 UI/Kg

n: 2837

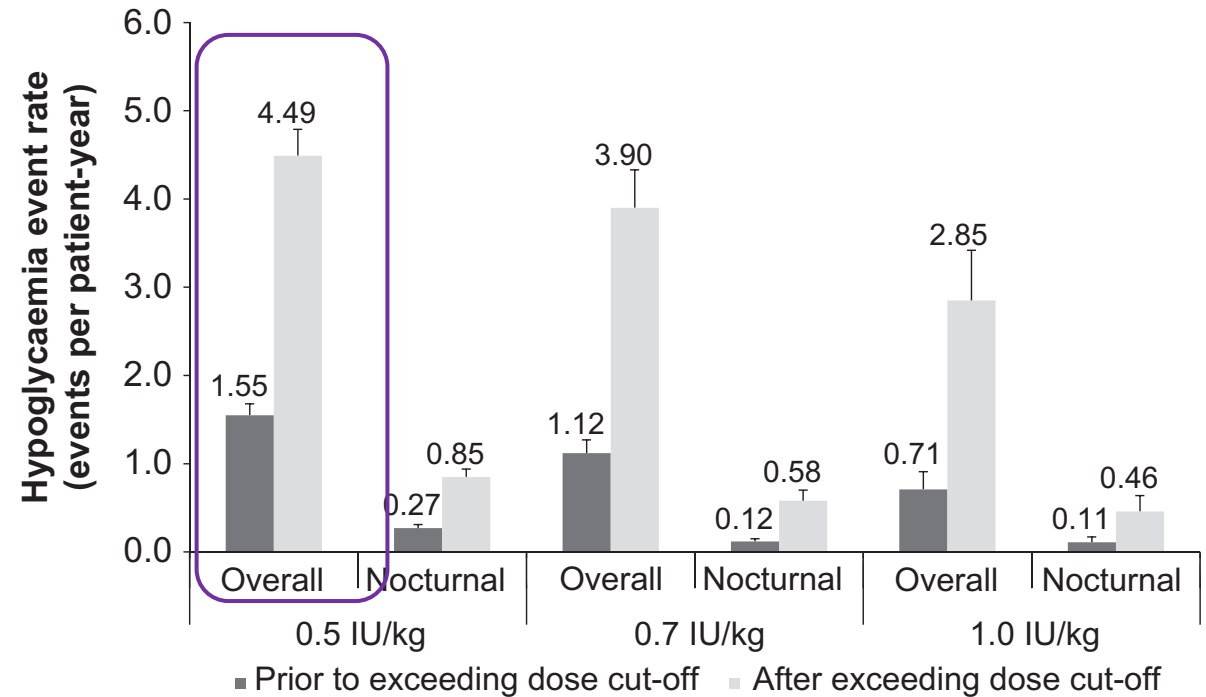
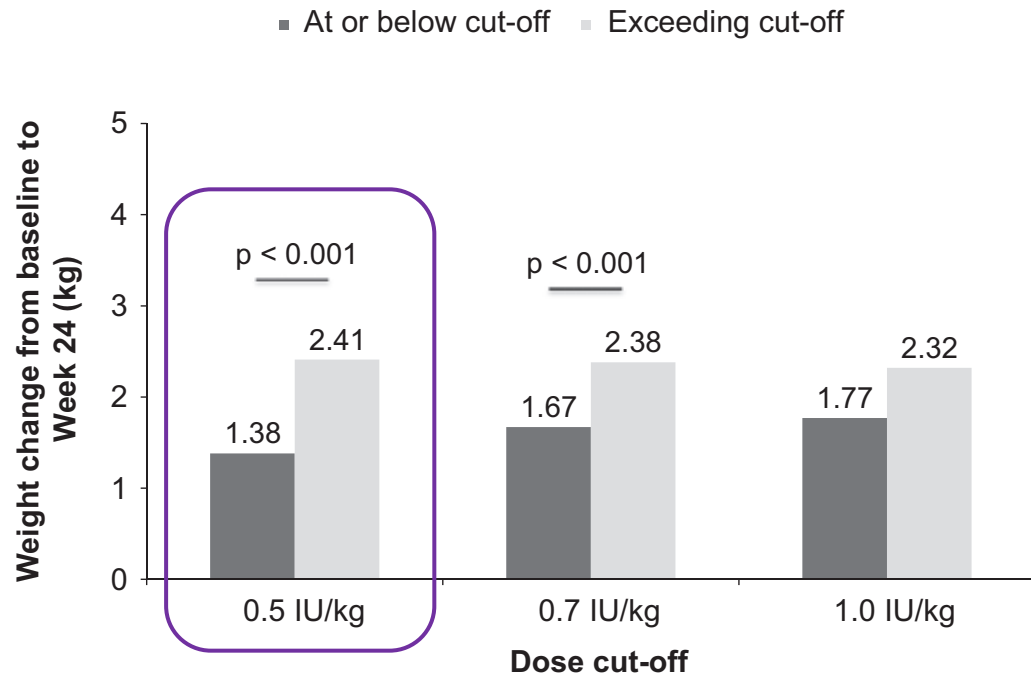
53-59 años

7.6 – 9.5 años con DM2



# How much is too much? Outcomes in patients using high-dose insulin glargine

T. Reid,<sup>1</sup> L. Gao,<sup>2</sup> J. Gill,<sup>3</sup> A. Stuhr,<sup>3</sup> L. Traylor,<sup>3</sup> A. Vlajnic,<sup>3</sup> A. Rhinehart<sup>4</sup>





INSULINA



PESO

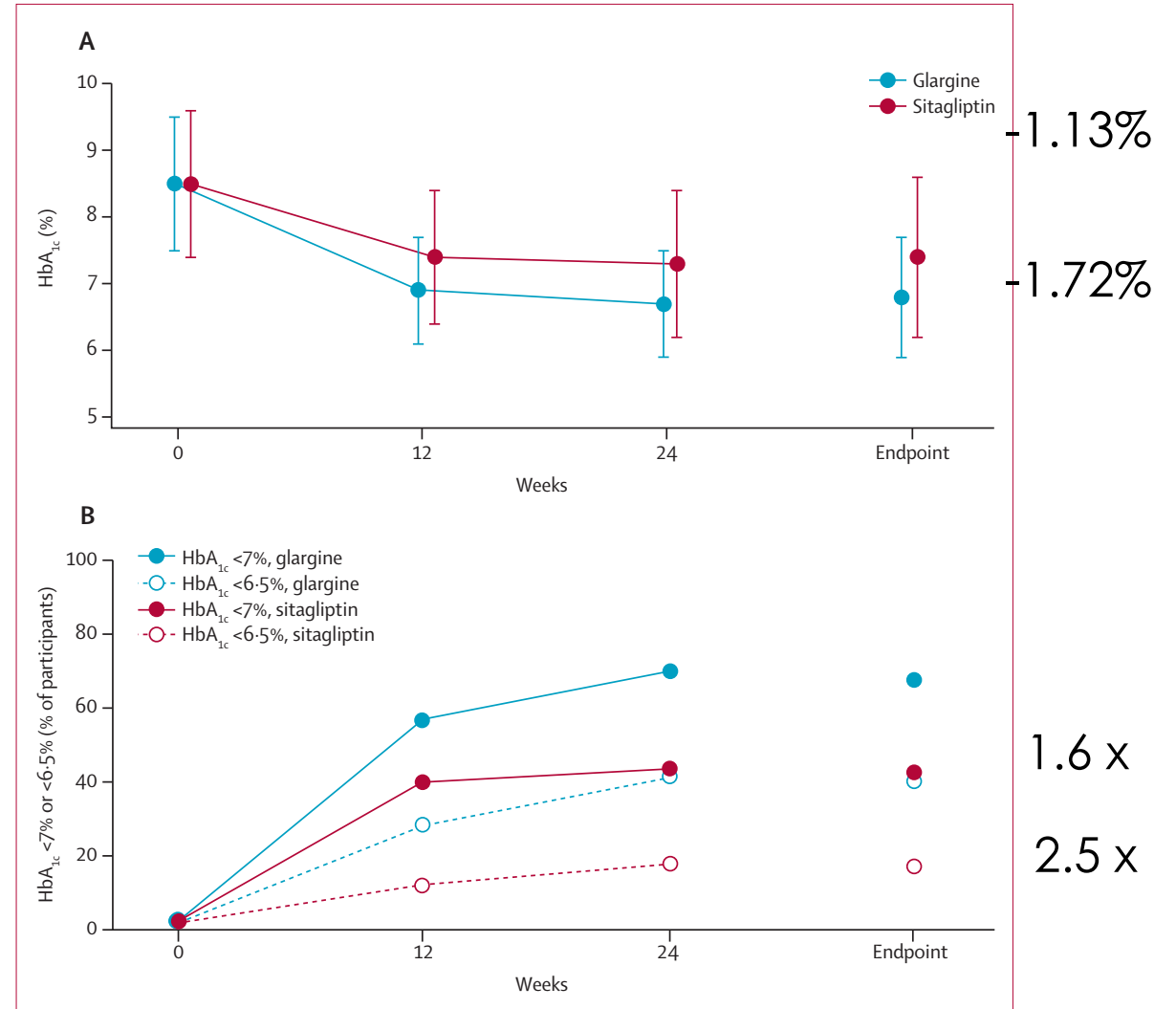




**Insulin glargine versus sitagliptin in insulin-naive patients with type 2 diabetes mellitus uncontrolled on metformin (EASIE): a multicentre, randomised open-label trial**



*Pablo Aschner\*, Juliana Chan\*, David R Owens, Sylvie Picard, Edward Wang, Marie-Paule Dain, Valérie Pilorget, Akram Echtaï, Vivian Fonseca, on behalf of the EASIE investigators*





# 0.5 x 0.5

“El control glucemico se alcanzó con 0.5 U/Kg y con tan solo una ganancia de peso de 0.5 Kg”

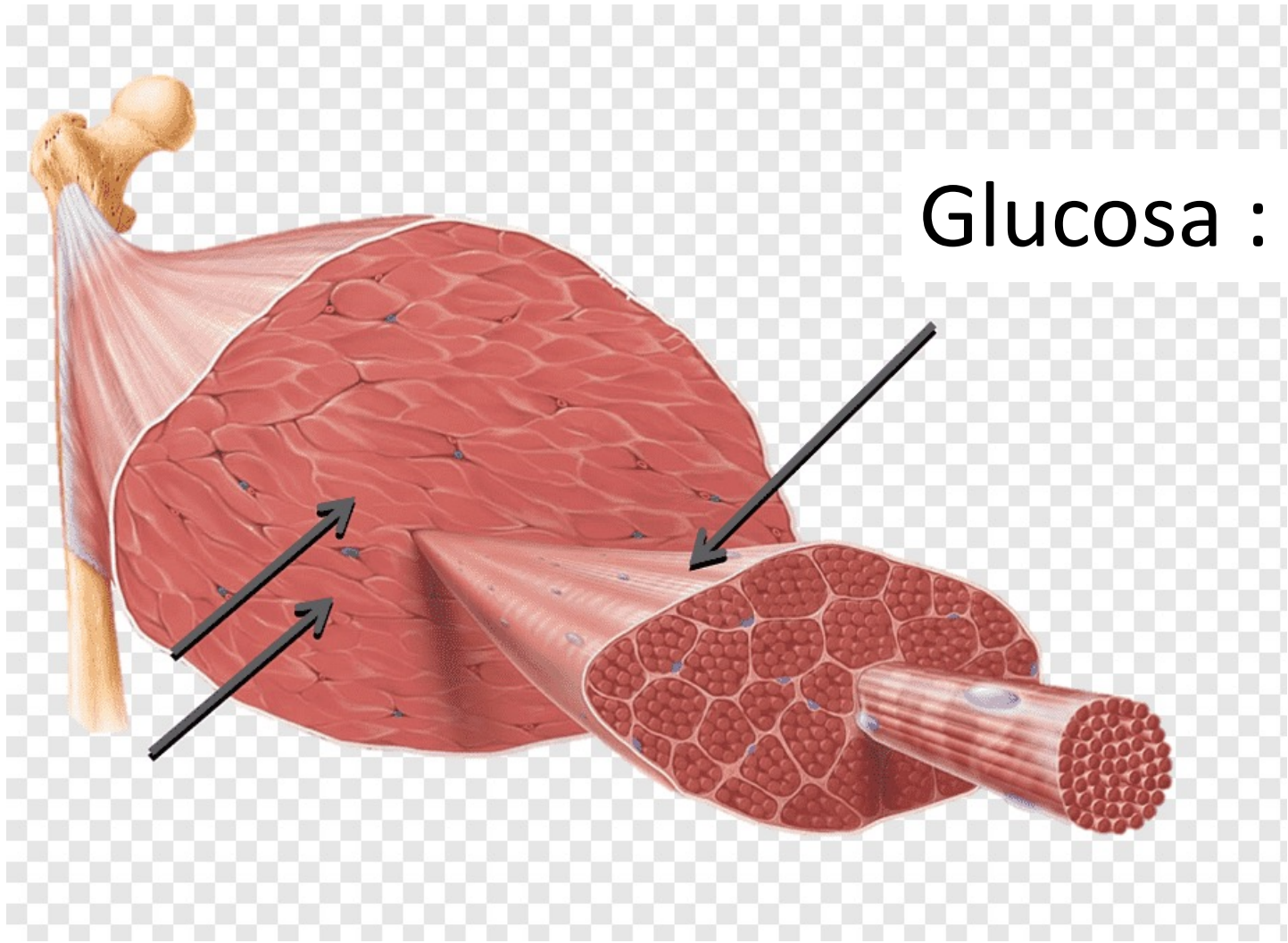




• **Lo que deberíamos aprender de la Insulina**



@eldoctorcastillo



Glucosa : 80%

# The Effect of Graded Doses of Insulin on Total Glucose Uptake, Glucose Oxidation, and Glucose Storage in Man

DANIEL THIEBAUD, ERIC JACOT, RALPH A. DEFRONZO, EVELYNE MAEDER, ERIC JEQUIER, AND JEAN-PIERRE FELBER

DIABETES, VOL. 31, NOVEMBER 1982



80%

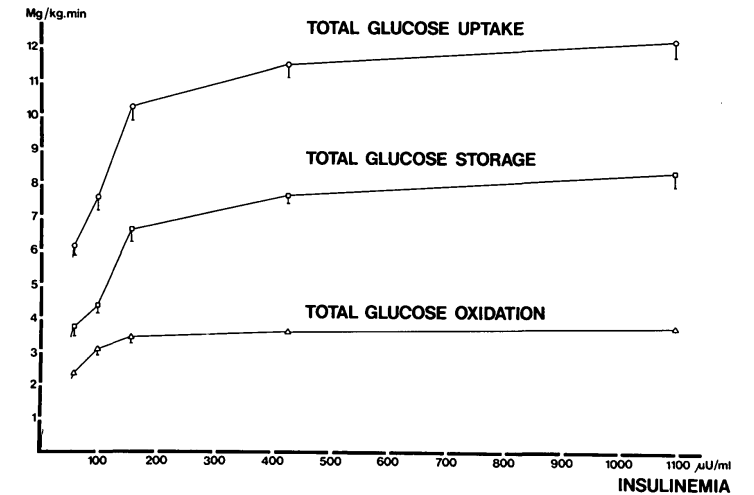
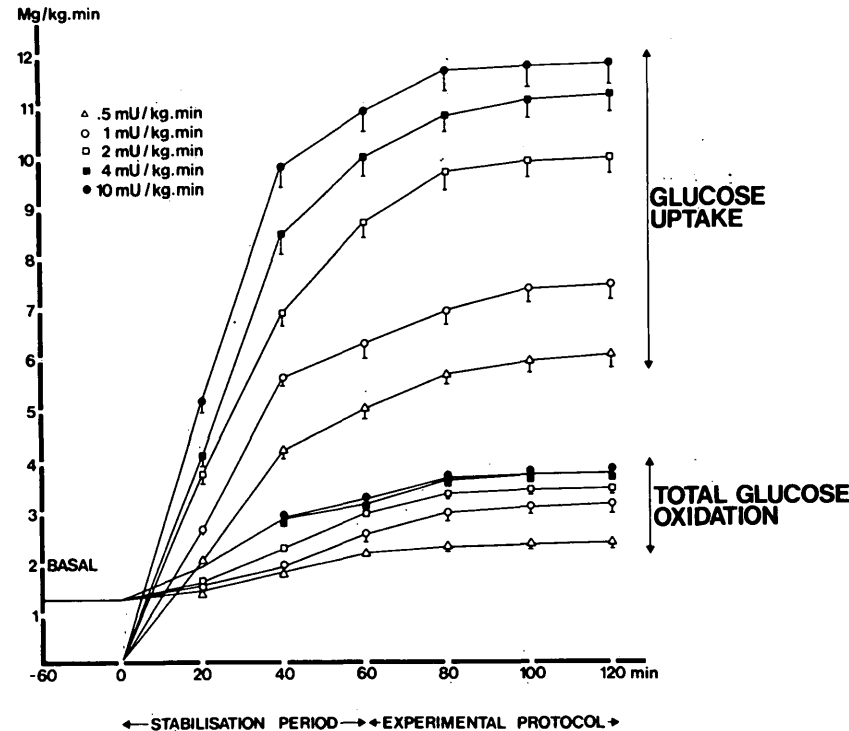
**Lipid** stores are large and potentially inexhaustible.

## Carbohydrate

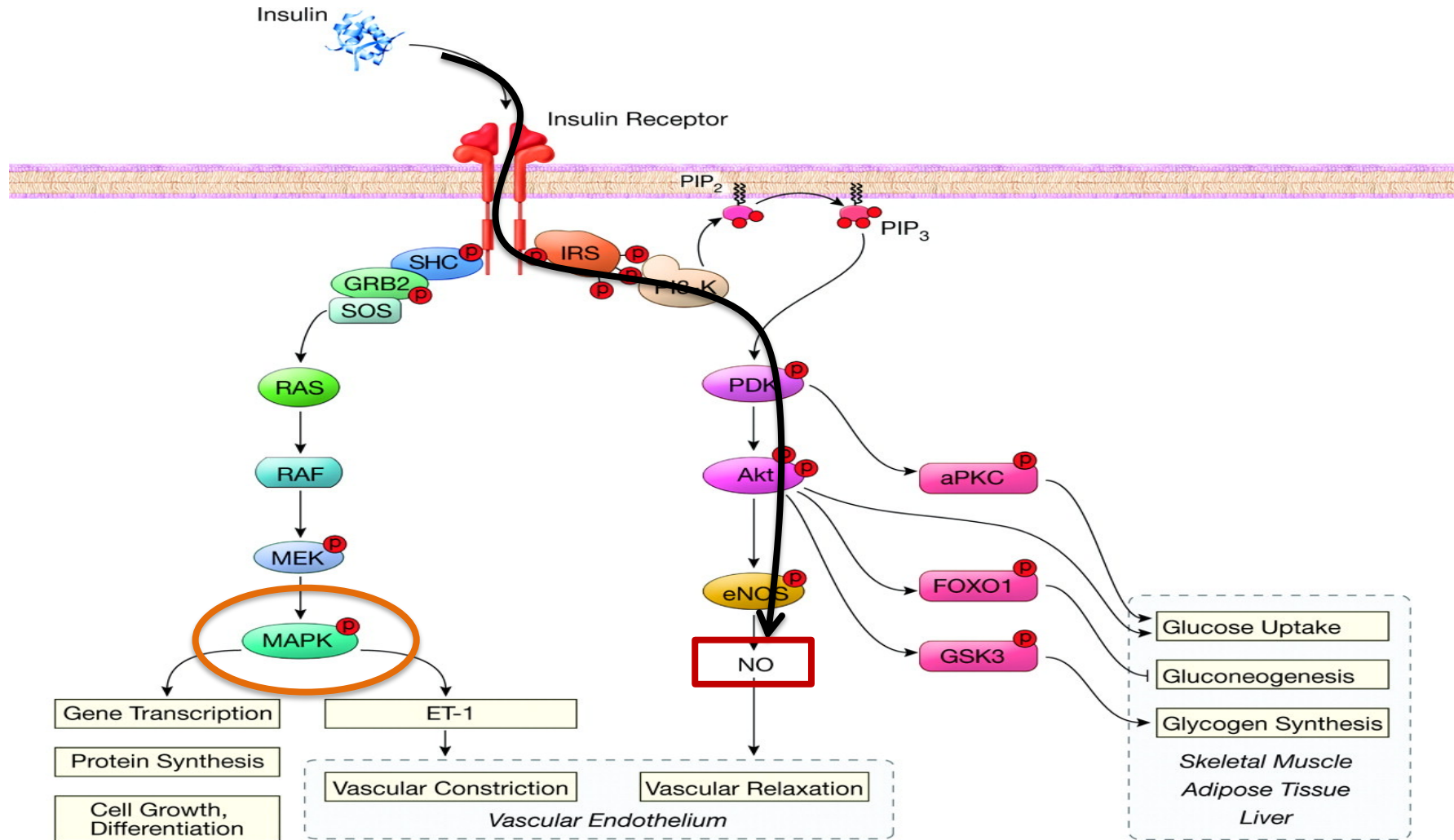
**Skeletal muscle**, 300–500 g of glycogen in

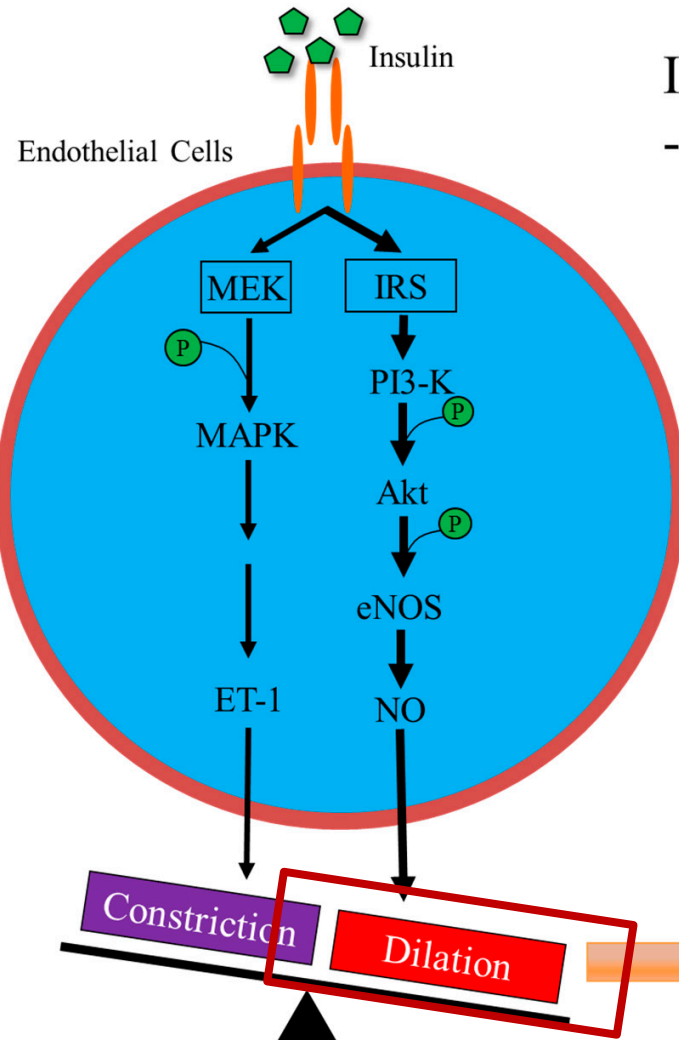
**Liver** 60–100 g of glycogen

**Glucose** 4–5 g of glucose circulating in the

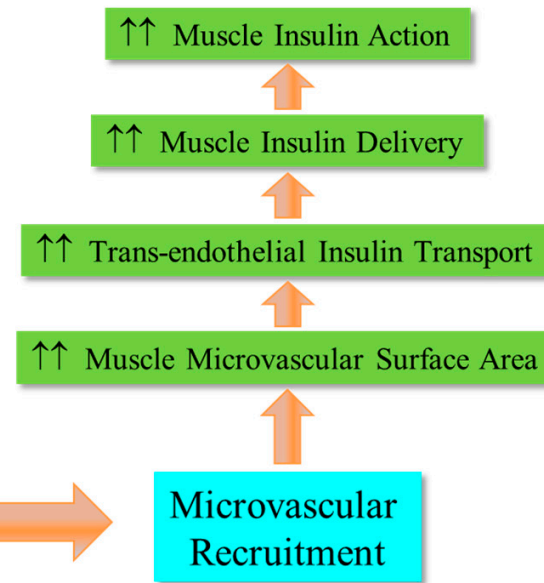


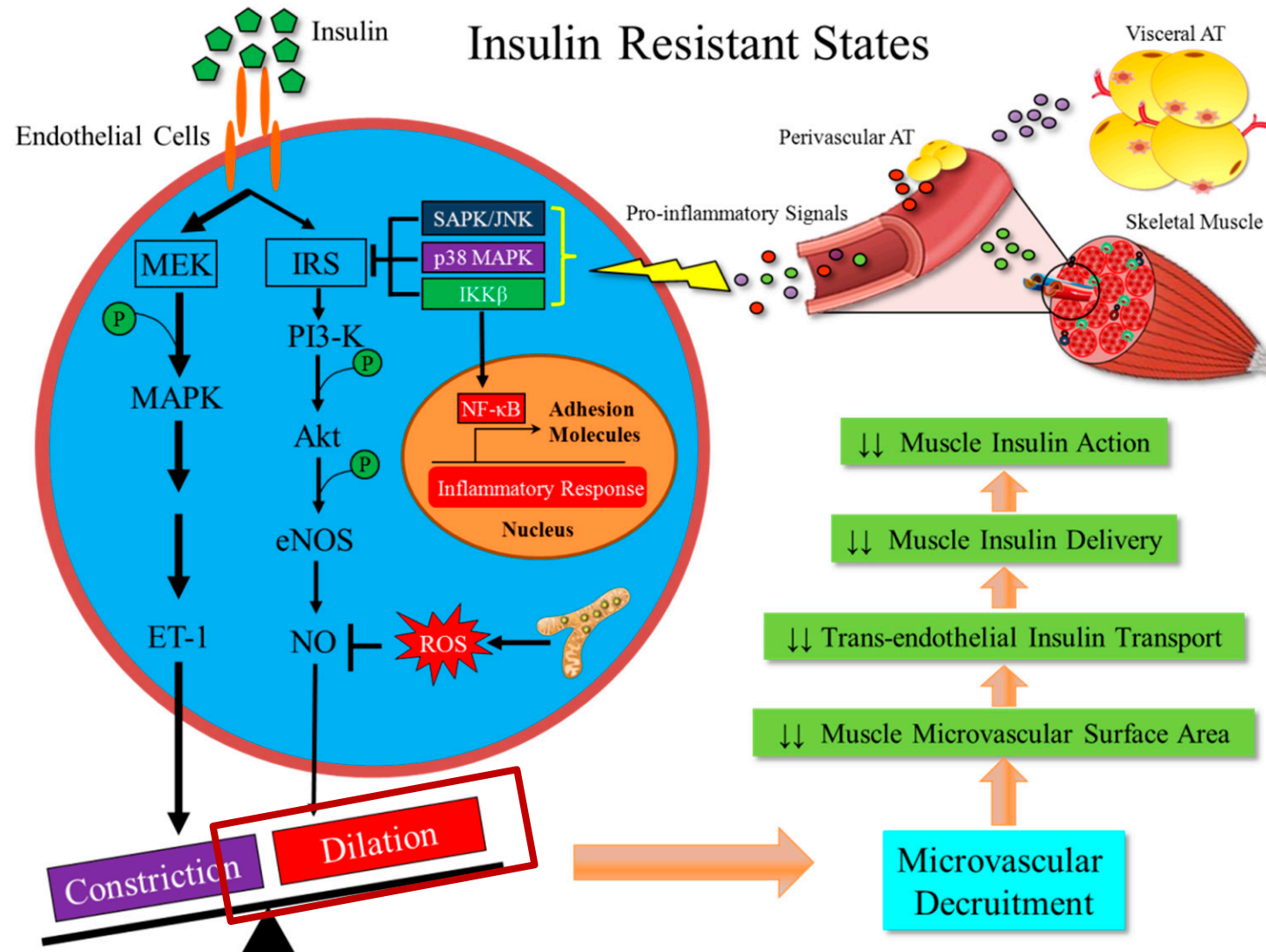
# General features of insulin signal transduction pathways





## Insulin Delivery and Action - A Feed Forward System





**Review**

*Am J Physiol Endocrinol Metab* 295: E732–E750, 2008.  
First published July 8, 2008; doi:10.1152/ajpendo.90477.2008.

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**HISTORICAL PERSPECTIVE |**

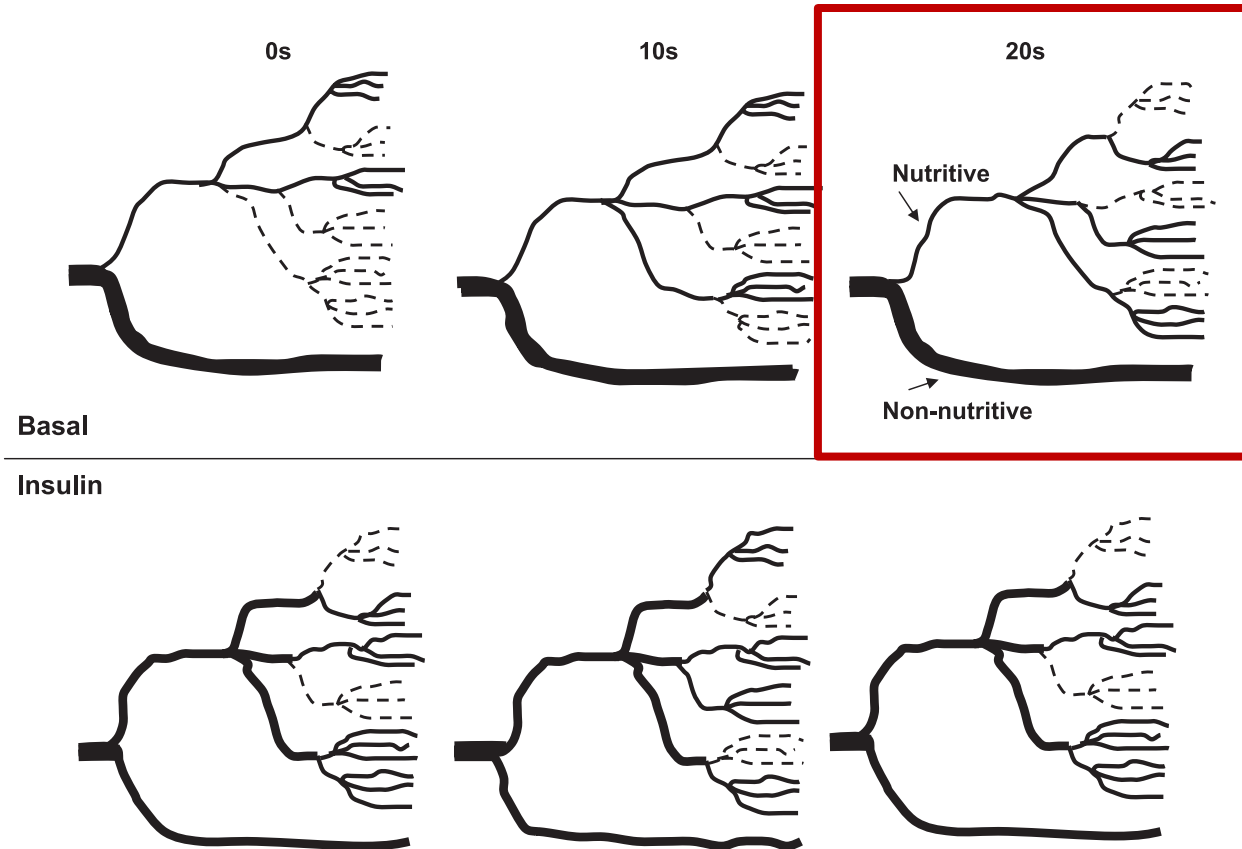
## Impaired microvascular perfusion: a consequence of vascular dysfunction and a potential cause of insulin resistance in muscle

**Michael G. Clark**

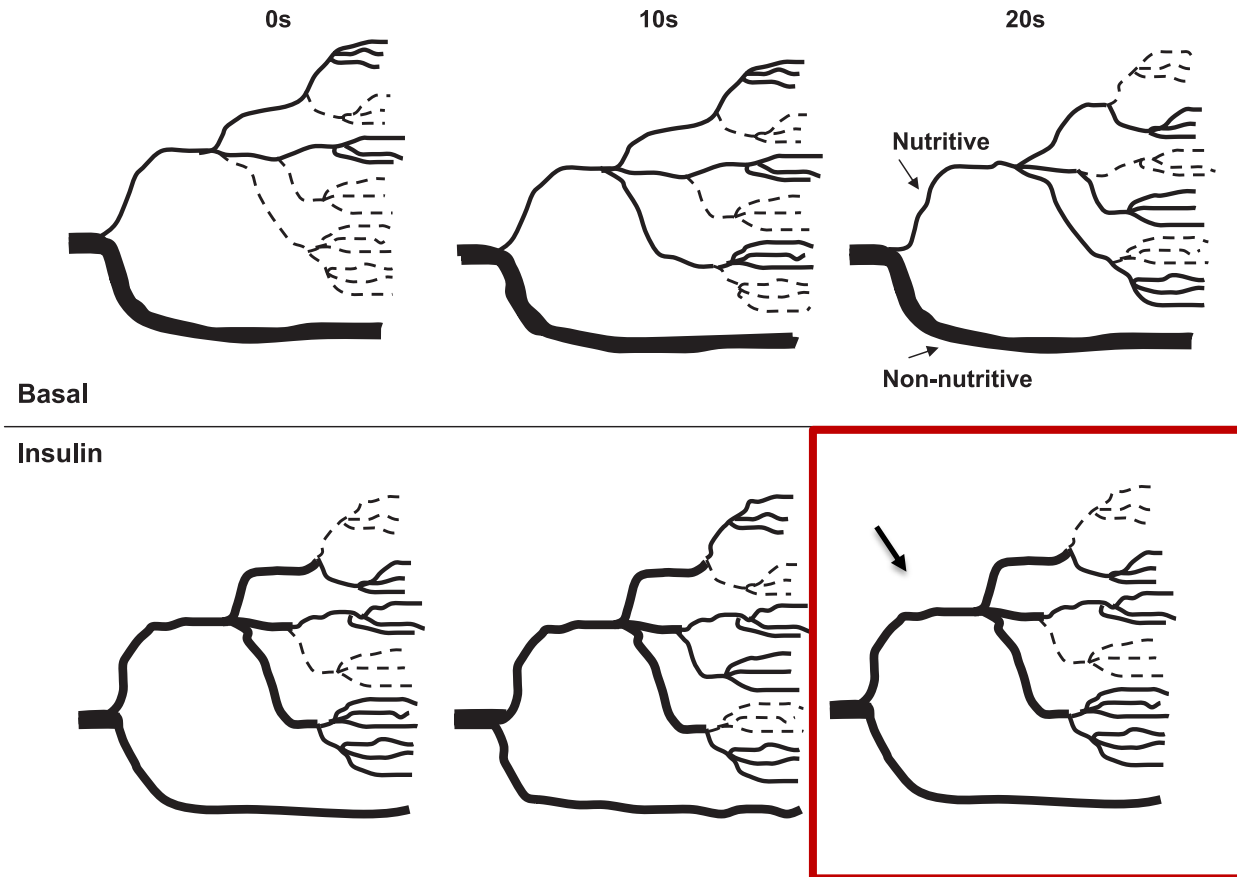
*Menzies Research Institute, University of Tasmania, Hobart, Australia*

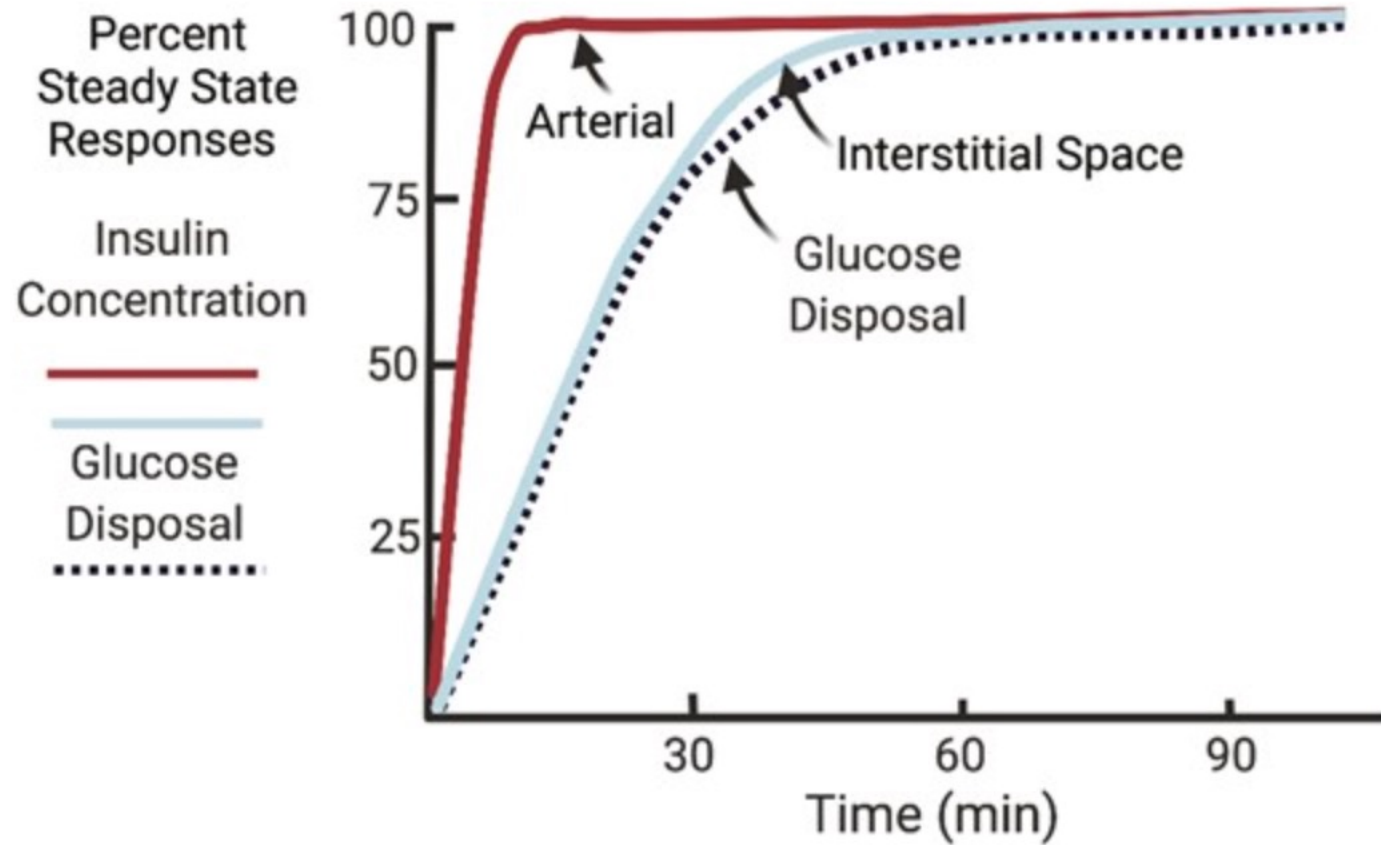
Submitted 29 May 2008; accepted in final form 7 July 2008

**Fig. 1. Proposed schematic blood flow patterns in muscle in vivo under basal conditions and following a physiological rise in plasma insulin**



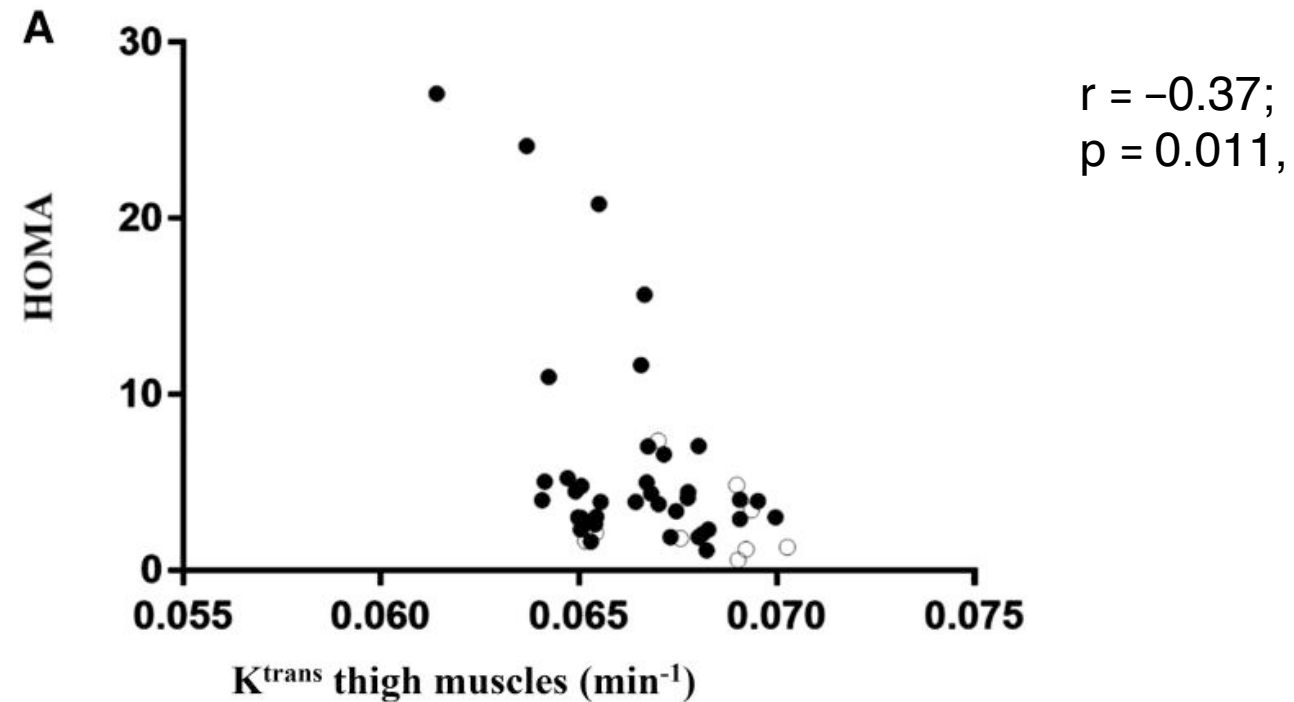
**Fig. 1. Proposed schematic blood flow patterns in muscle in vivo under basal conditions and following a physiological rise in plasma insulin**





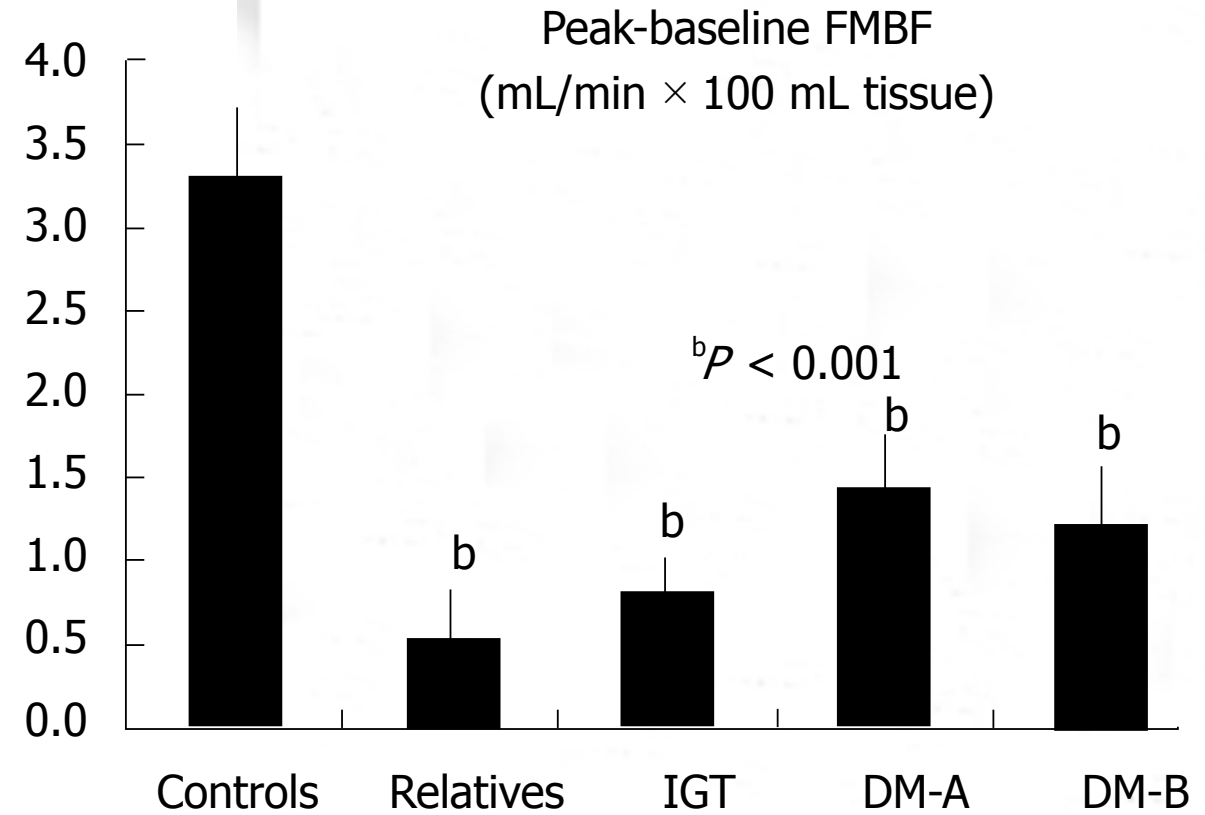
## Insulin Resistance Is Associated With Reduced Capillary Permeability of Thigh Muscles in Patients With Type 2 Diabetes

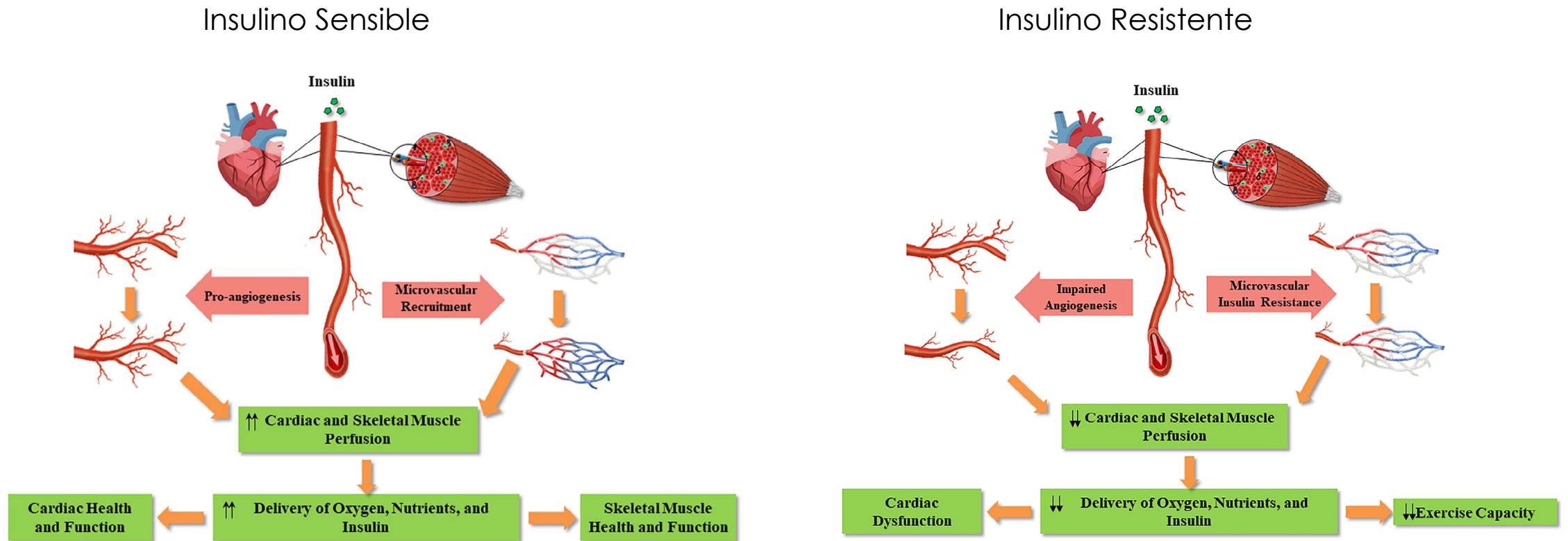

Correlation analysis of the thigh muscles' constant of **permeability** ( $K^{\text{trans}}$ ) with HOMA index

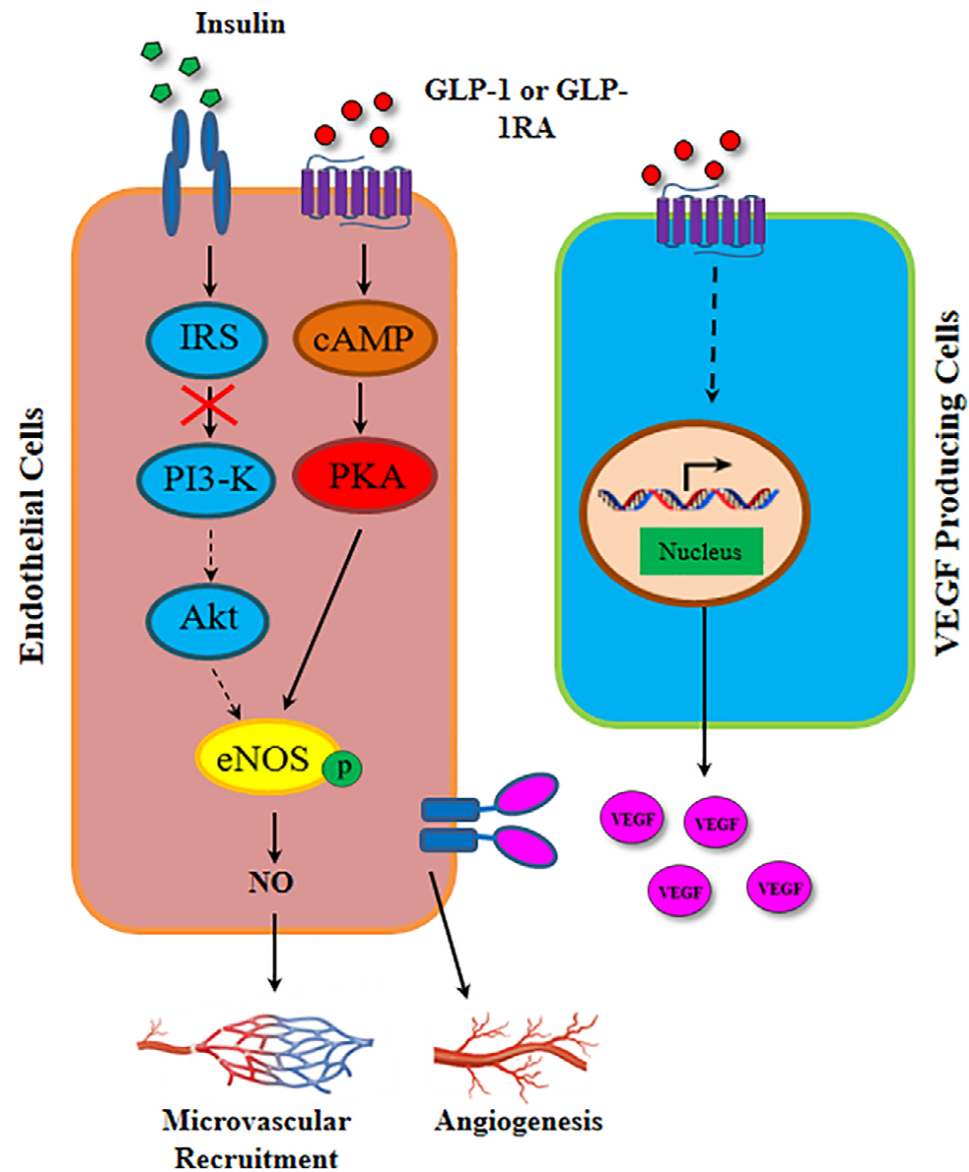



## Insulin Resistance Is Associated With Reduced Capillary Permeability of Thigh Muscles in Patients With Type 2 Diabetes

Forearm muscle blood flow peak-baseline



**GLP-1 and insulin regulation of skeletal and cardiac muscle microvascular perfusion in type 2 diabetes**Kaitlin M. Love<sup>1</sup> | Jia Liu<sup>1</sup> | Judith G. Regensteiner<sup>2,3</sup> | Jane E.B. Reusch<sup>2,3,4</sup> | Zhenqi Liu<sup>1</sup> 

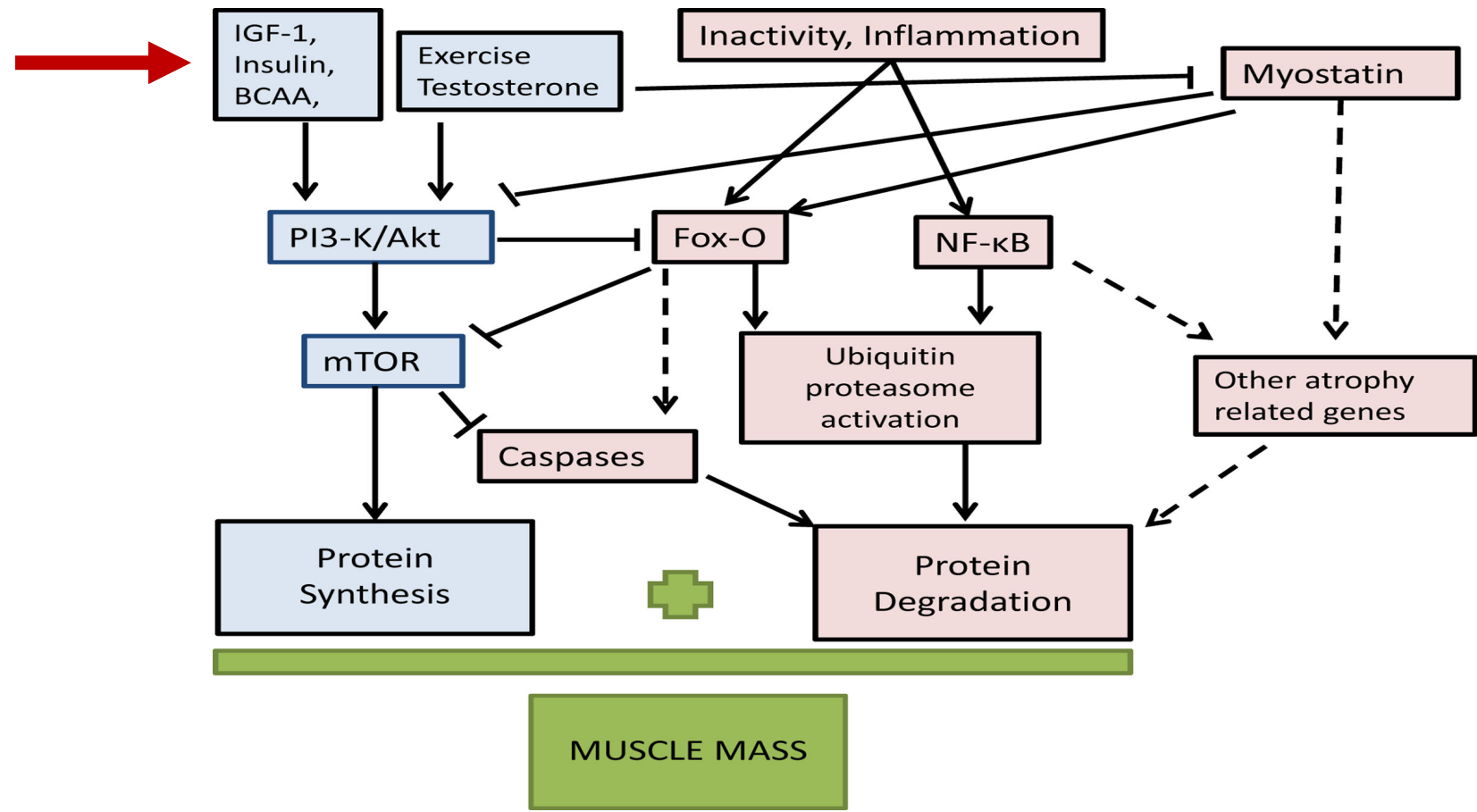
**GLP-1 and insulin regulation of skeletal and cardiac muscle microvascular perfusion in type 2 diabetes**Kaitlin M. Love<sup>1</sup> | Jia Liu<sup>1</sup> | Judith G. Regensteiner<sup>2,3</sup> | Jane E.B. Reusch<sup>2,3,4</sup> | Zhenqi Liu<sup>1</sup> 

## Insulin Glargine is More Suitable Than Exenatide in Preventing Muscle Loss in Non-Obese Type 2 Diabetic Patients with NAFLD

► **Table 2** Interaction between changes in the Psoas Muscle Area (PMA) among subgroups.

	Unadjusted			Adjusted *		
	Changes of PMA ( mm <sup>2</sup> )	P value	Interac- tion test P value	Changes of PMA (mm <sup>2</sup> )	P value	Interac- tion test P value
	β (95 %CI)			β (95 %CI)		
Insulin Glargine vs. Exenatide						
Total	178.54 (−99.98, 457.07)	0.214		202.39 (−76.34, 481.12)	0.16	
Male	298.92 (−214.97, 812.81)	0.263	0.362	368.48 (−128.99, 865.95)	0.158	0.189
Female	44.91 (−123.45, 213.27)	0.605		11.22 (−169.41, 191.84)	0.904	
Age <60y	208.73 (−133.59, 551.06)	0.238	0.758	185.43 (−131.85, 502.72)	0.258	0.963
Age ≥60y	99.04 (−164.34, 362.42)	0.477		144.83 (−114.57, 404.23)	0.316	
BMI <28 kg/m <sup>2</sup>	403.04 (−17.43, 823.51)	0.069	0.038	560.64 (77.88, 1043.40)	0.031	0.009
BMI ≥28 kg/m <sup>2</sup>	−112.92 (−466.65, 240.80)	0.537		−114.21 (−467.10, 238.68)	0.532	

\* adjusted for baseline PMA, age, sex and center, if appropriate; PMA: posas muscle area; BMI: body mass index.



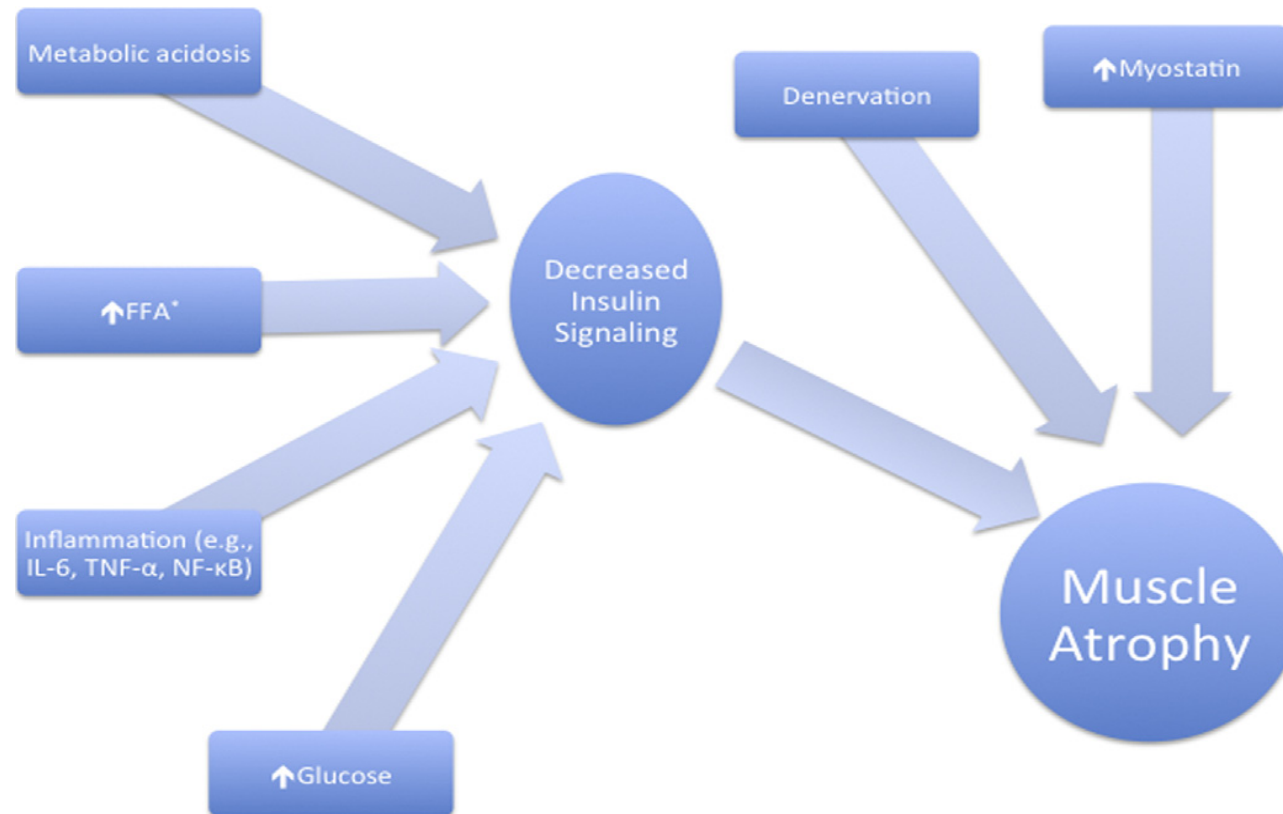


Review

The regulation of muscle protein turnover in diabetes<sup>☆</sup>

Biruh Workeneh, Mandeep Bajaj\*

Baylor College of Medicine, 1709 Dryden, Suite 900, Houston, TX 77030, USA

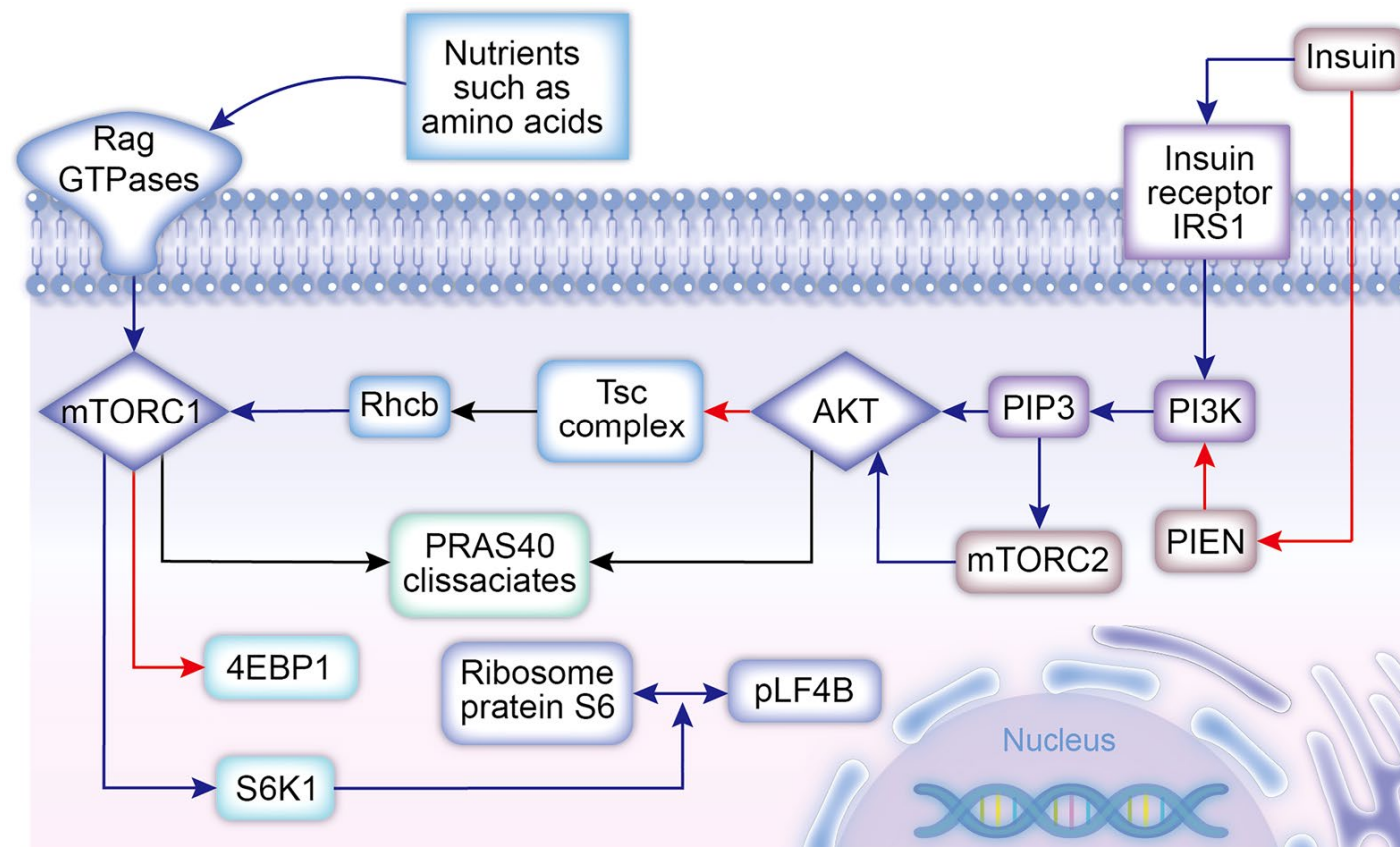


**Fig. 1.** This figure summarizes the major stimuli for muscle atrophy in type 2 diabetes. \*FFA, free fatty acids.

## REVIEW

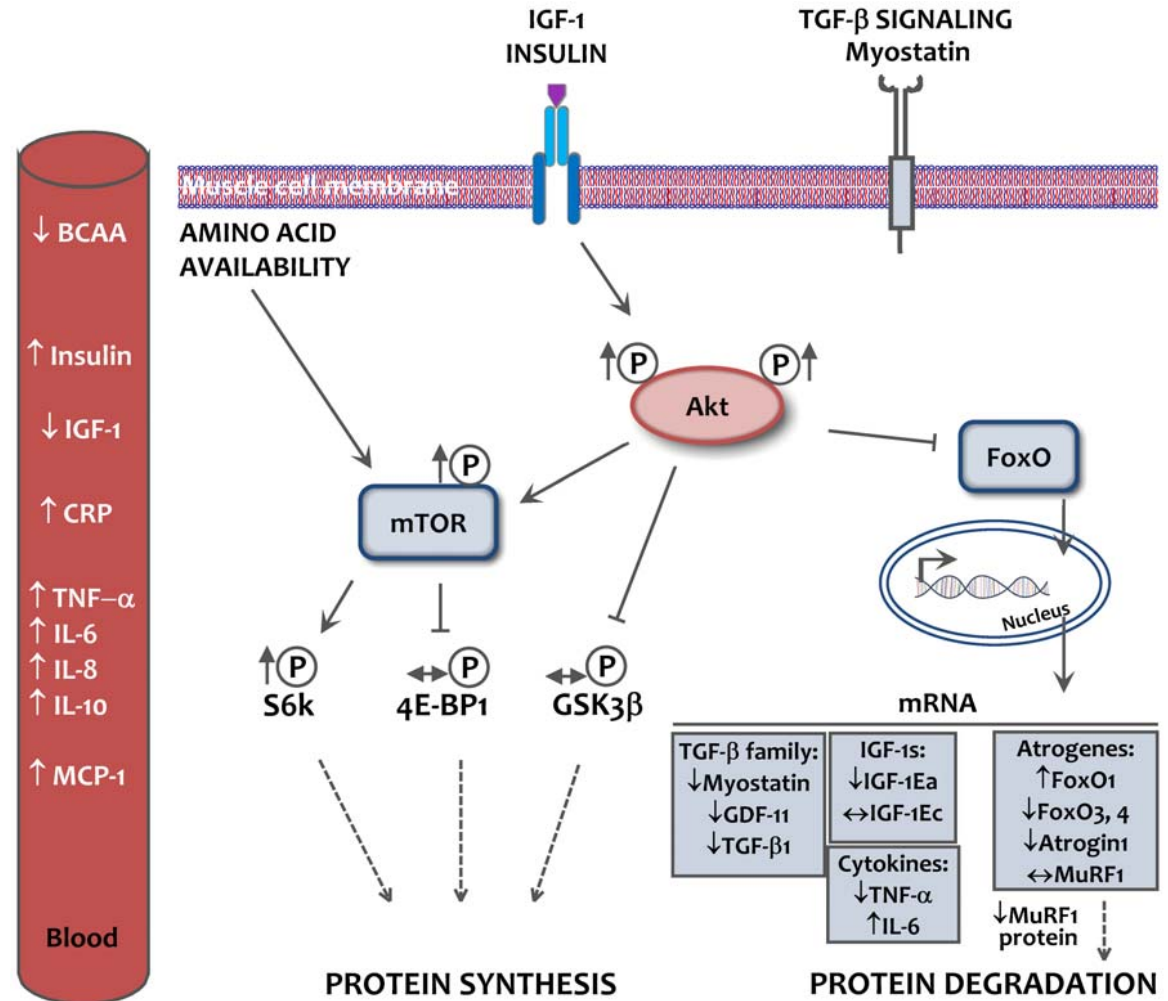
## Open Access

## Causal relationship between insulin resistance and sarcopenia

Zi-jian Liu<sup>1</sup> and Cui-feng Zhu<sup>2\*</sup>

# Activated Protein Synthesis and Suppressed Protein Breakdown Signaling in Skeletal Muscle of Critically Ill Patients

Jakob G. Jespersen<sup>1\*</sup>, Anders Nedergaard<sup>1</sup>, Søren Reitelseder<sup>1</sup>, Ulla R. Mikkelsen<sup>1</sup>, Kasper J. Dideriksen<sup>1</sup>, Jakob Agergaard<sup>1</sup>, Frederik Kreiner<sup>2</sup>, Frank C. Pott<sup>3</sup>, Peter Schjerling<sup>1</sup>, Michael Kjaer<sup>1</sup>



## Effect of insulin on human skeletal muscle protein synthesis is modulated by insulin-induced changes in muscle blood flow and amino acid availability

Satoshi Fujita<sup>1,3</sup>, Blake B. Rasmussen<sup>2,4</sup>, Jerson G. Cadenas<sup>1,3</sup>, James J. Grady<sup>5</sup>, and Elena Volpi<sup>1,3</sup>

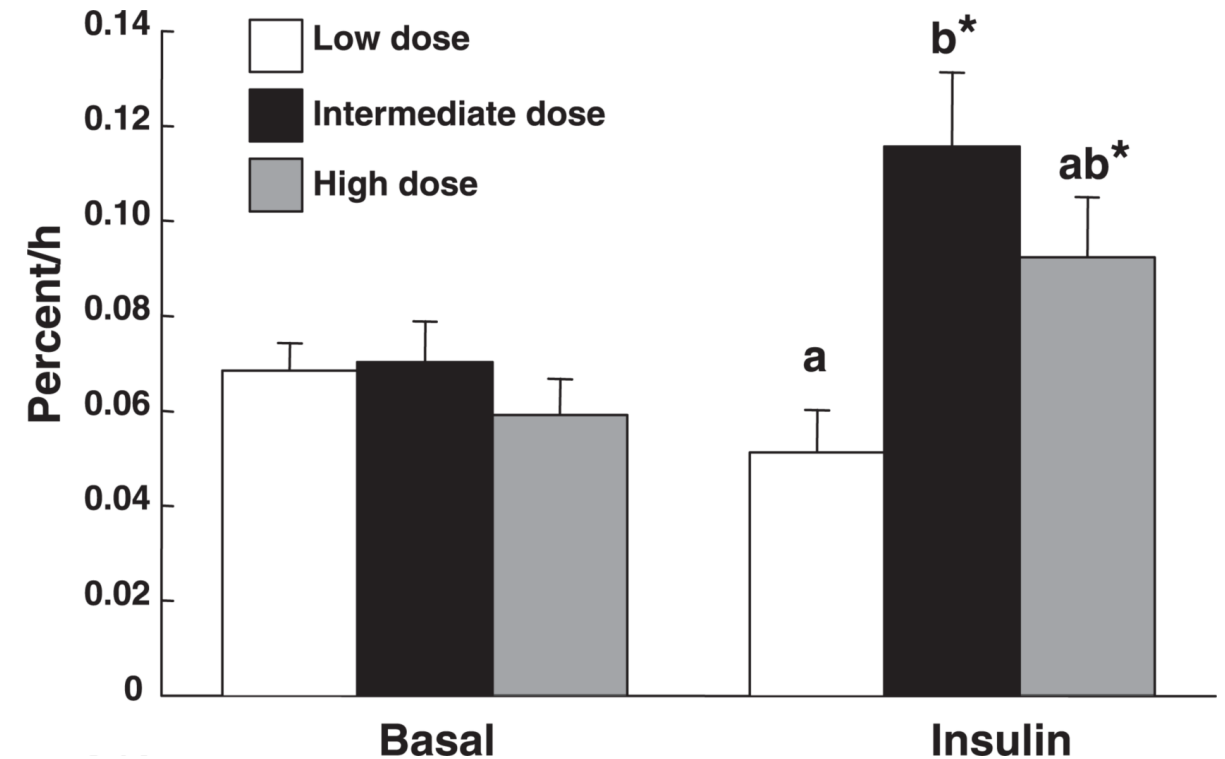
Table 1

Physical characteristics of the subjects

	LD	ID	HD	<i>P</i>
<i>n</i>	6	7	6	
Sex	4 M 2 F	3 M 4 F	4 M 2 F	0.60
Age, yr	29±3	25±2	28±2	0.51
Height, cm	169±5	169±4	170±2	0.99
Body weight, kg	72±7	63±6	74±5	0.44
Body mass index, kg/m <sup>2</sup>	25±2	22±1	26±2	0.19
Fat-free mass, kg	50±4	45±5	54±3	0.30
Fat mass, kg	18±4	11±1	17±3	0.21
Leg volume, liters	9.5±1.1	8.4±5.9	9.9±3.5	0.33
Leg muscle mass, kg	8.6±8.9	7.3±8.9	9.9±3.3	0.21

Values are means ± SE. Subjects were randomized to 3 groups receiving a low (LD), intermediate (ID), or high dose (HD) insulin infusion. *P*, significance level for differences between groups.

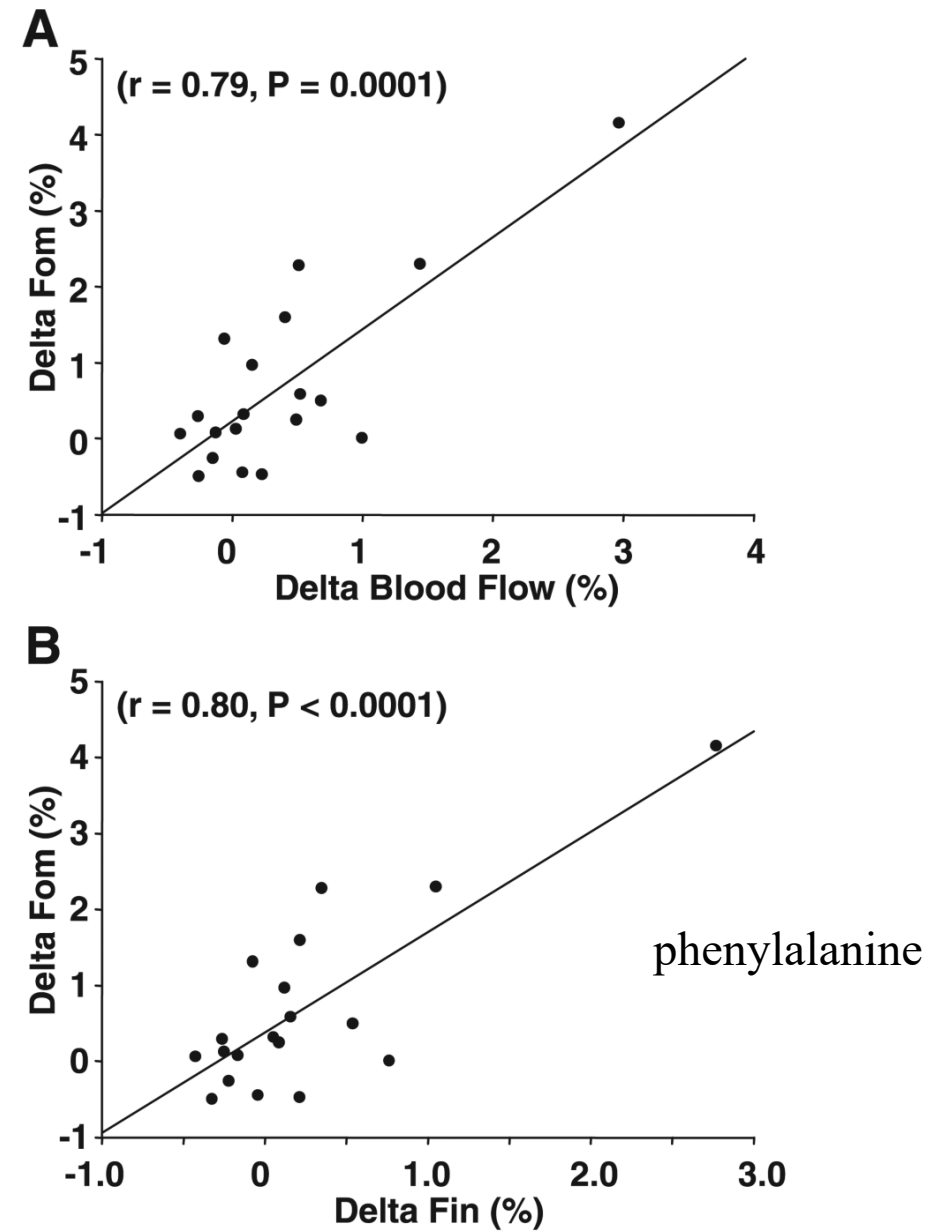
## Muscle protein fractional synthetic rates



**Effect of insulin on human skeletal muscle protein synthesis is modulated by insulin-induced changes in muscle blood flow and amino acid availability**

Satoshi Fujita<sup>1,3</sup>, Blake B. Rasmussen<sup>2,4</sup>, Jerson G. Cadenas<sup>1,3</sup>, James J. Grady<sup>5</sup>, and Elena Volpi<sup>1,3</sup>

Síntesis muscular



# Muscle Protein Synthesis (MPS)

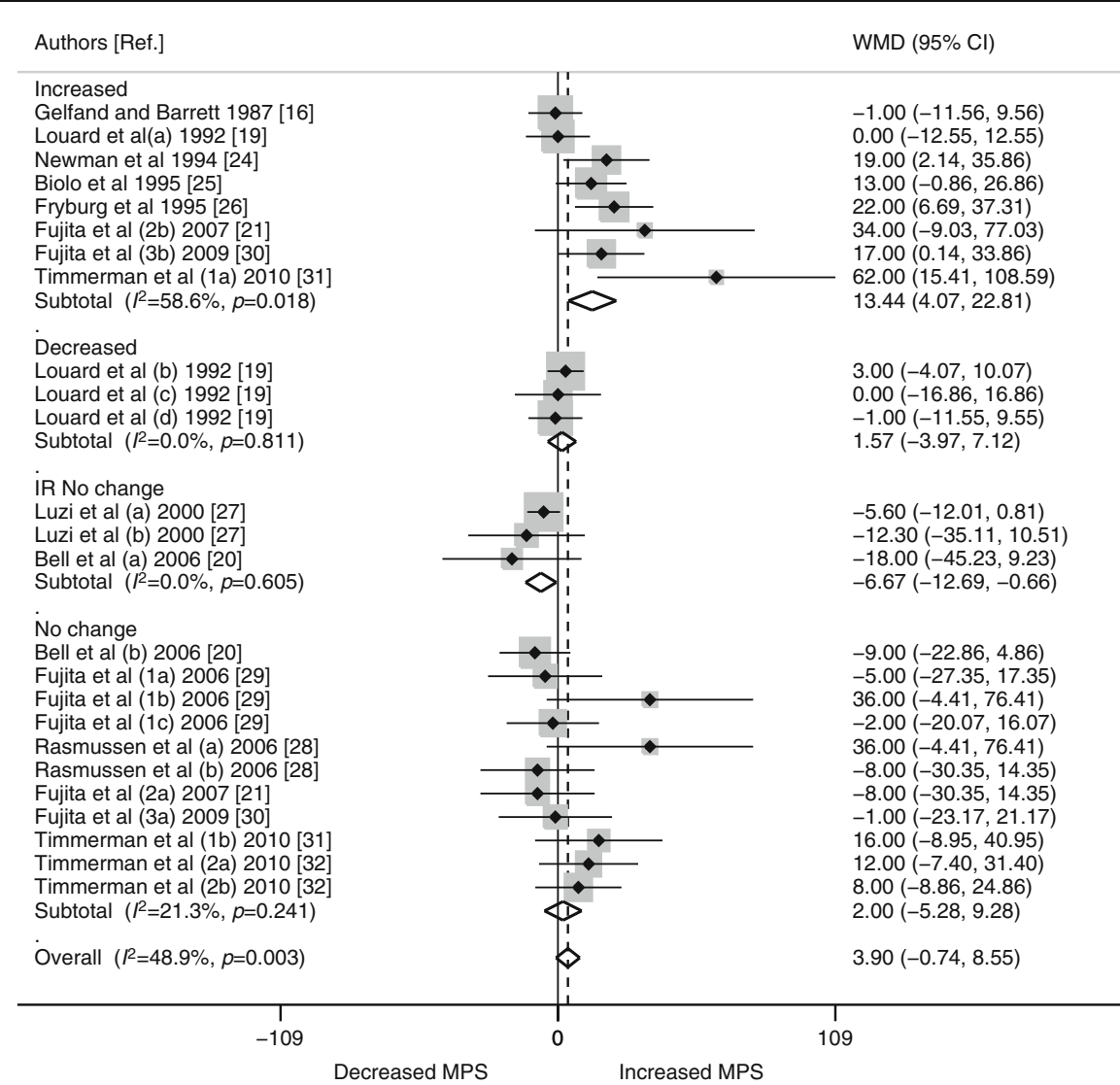
Diabetologia (2016) 59:44–55  
DOI 10.1007/s00125-015-3751-0



META-ANALYSIS

## Role of insulin in the regulation of human skeletal muscle protein synthesis and breakdown: a systematic review and meta-analysis

Haitham Abdulla<sup>1</sup> · Kenneth Smith<sup>1</sup> · Philip J. Atherton<sup>1</sup> · Iskandar Idris<sup>1</sup>



# Muscle Protein Synthesis (MPS)

Con disponibilidad de a.a.

Diabetologia (2016) 59:44–55  
DOI 10.1007/s00125-015-3751-0

META-ANALYSIS

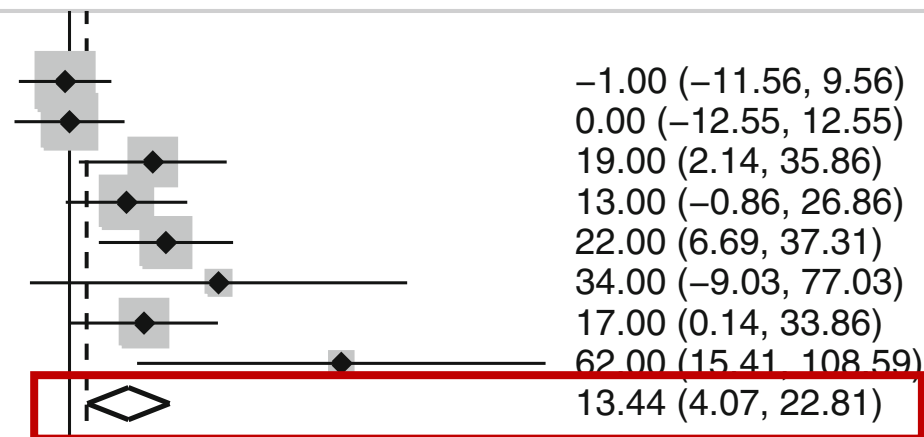
## Role of insulin in the regulation of hum synthesis and breakdown: a systematic

Haitham Abdulla<sup>1</sup> · Kenneth Smith<sup>1</sup> · Philip J. Atherton<sup>1</sup> · Iskandar

Authors [Ref.]

WMD (95% CI)

Increased  
 Gelfand and Barrett 1987 [16]  
 Louard et al(a) 1992 [19]  
 Newman et al 1994 [24]  
 Biolo et al 1995 [25]  
 Fryburg et al 1995 [26]  
 Fujita et al (2b) 2007 [21]  
 Fujita et al (3b) 2009 [30]  
 Timmerman et al (1a) 2010 [31]  
 Subtotal ( $I^2=58.6%$ ,  $p=0.018$ )



# Muscle Protein Break- down (MPB)

Diabetologia (2016) 59:44–55  
DOI 10.1007/s00125-015-3751-0



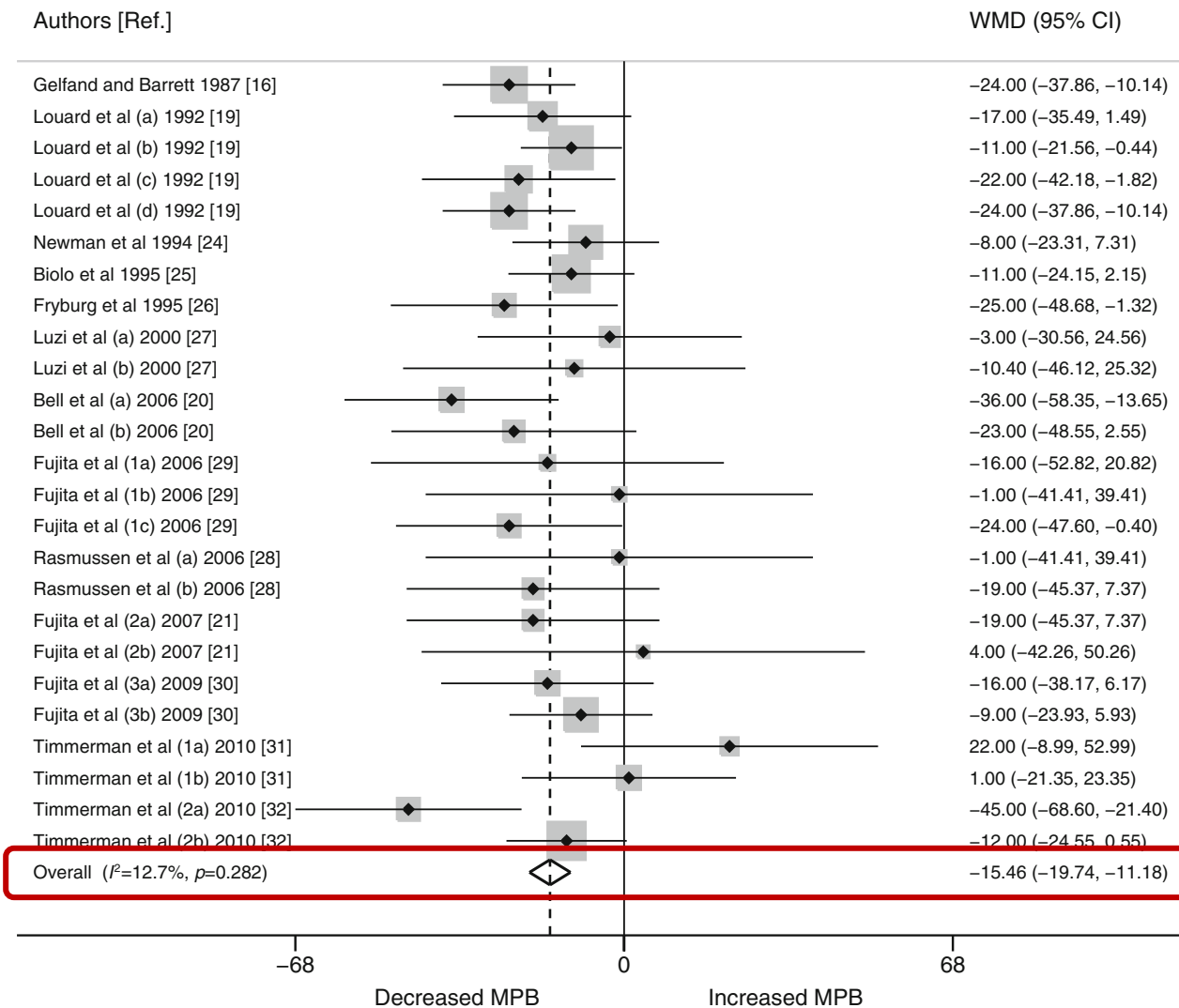
META-ANALYSIS

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-15.46 nmol/100 ml volumen de pierna/min

-15.46 nmol menos de aminoácidos que salen de cada 100 ml de musculo, cada miniuto



## Muscle Protein Break- down (MPB)

Diabetologia (2016) 59:44–55  
DOI 10.1007/s00125-015-3751-0



META-ANALYSIS

### Role of insulin in the regulation of human skeletal muscle protein synthesis and breakdown: a systematic review and meta-analysis

Haitham Abdulla<sup>1</sup> · Kenneth Smith<sup>1</sup> · Philip J. Atherton<sup>1</sup> · Iskandar Idris<sup>1</sup>

- Sin insulina:  
→ el músculo “pierde” aminoácidos constantemente
- Con insulina:  
→ esa fuga baja en ~15–20%

👉 No construyes más músculo, pero **pierdes menos cada minuto**

-15.46 nmol/100 ml volumen de pierna/min

# Muscle Protein Break- down (MPB)

Diabetologia (2016) 59:44–55  
DOI 10.1007/s00125-015-3751-0



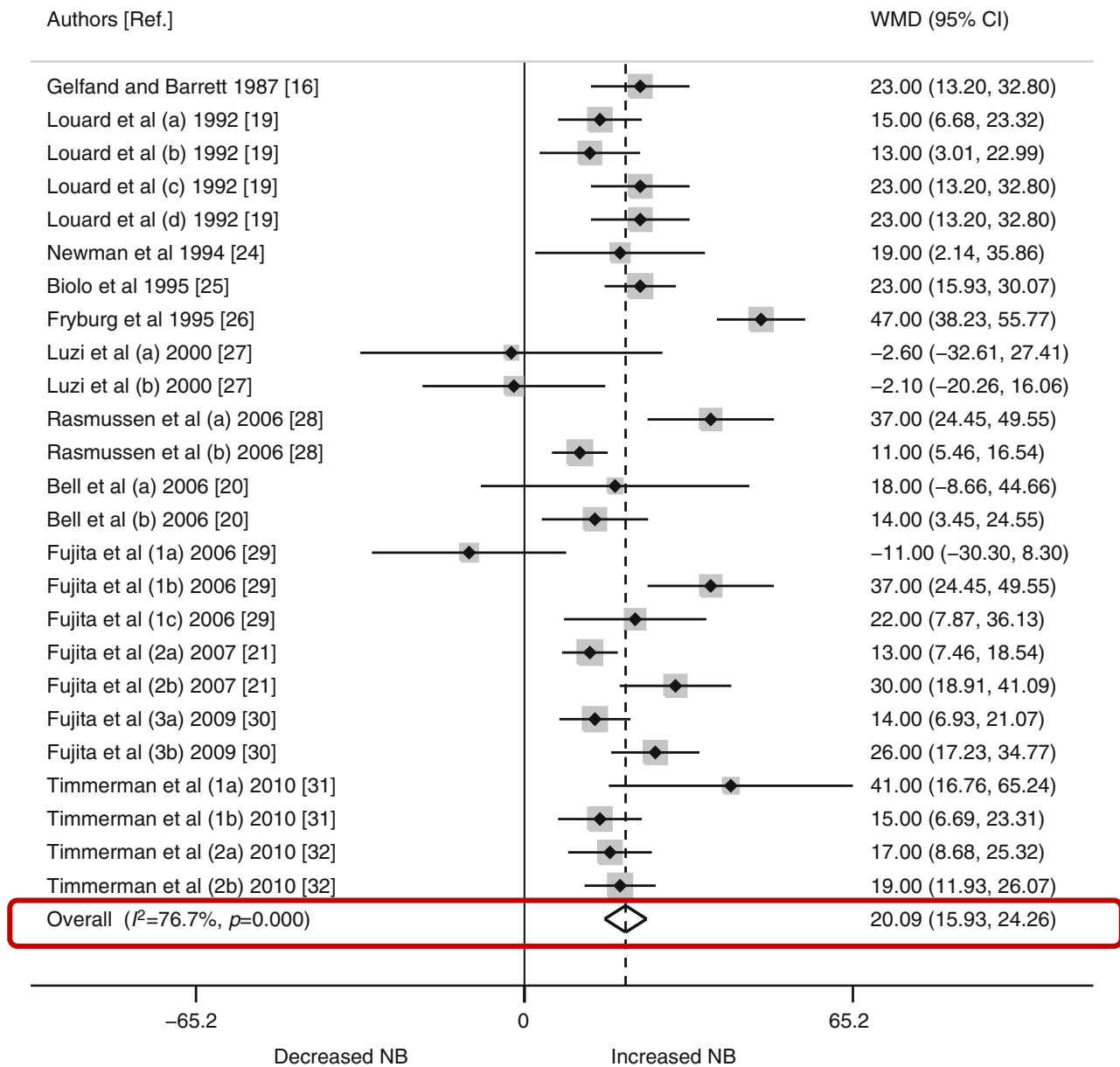
META-ANALYSIS

## Role of insulin in the regulation of human skeletal muscle protein synthesis and breakdown: a systematic review and meta-analysis

Haitham Abdulla<sup>1</sup> · Kenneth Smith<sup>1</sup> · Philip J. Atherton<sup>1</sup> · Iskandar Idris<sup>1</sup>

A pesar de su efecto variable sobre la síntesis, la insulina aumenta significativamente el balance neto de proteínas musculares

20.09 nmol/100 ml volumen de pierna/min

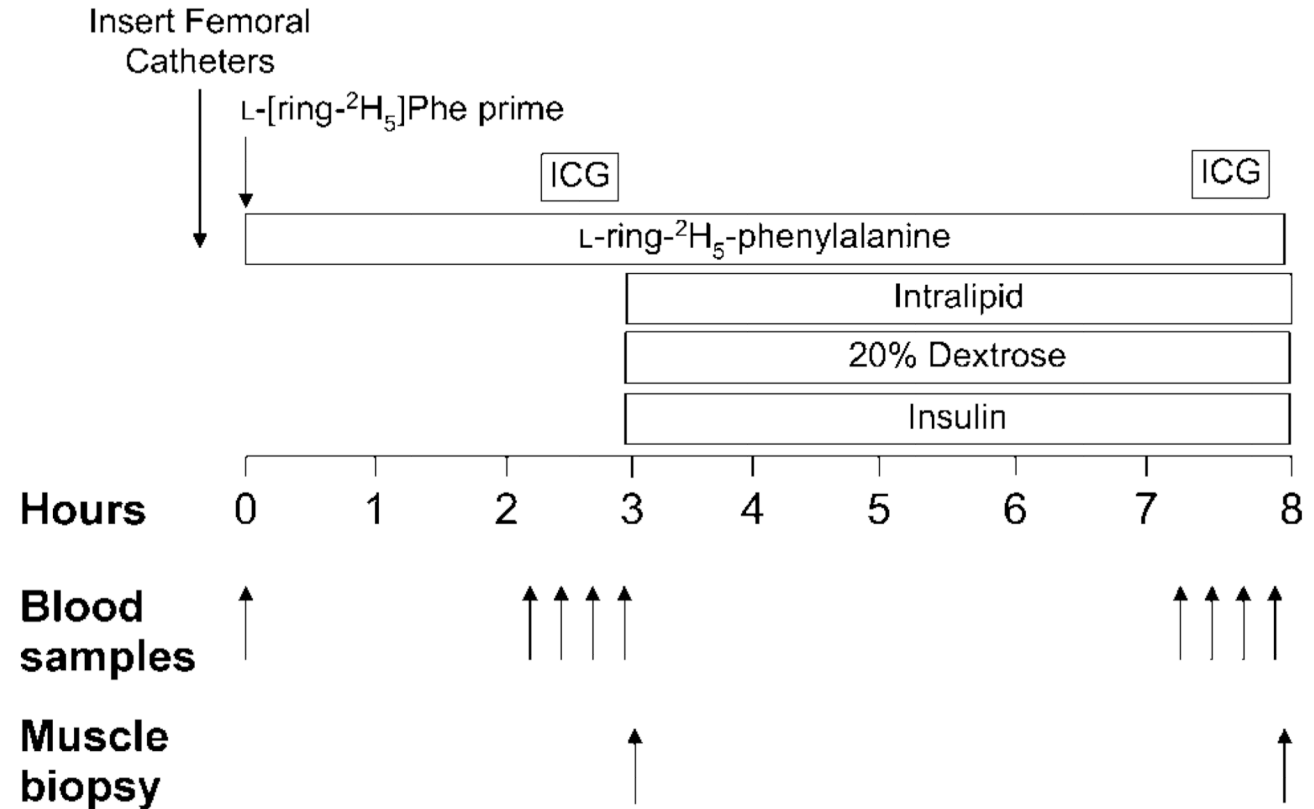




Estudios Clínicos

## Skeletal Muscle Protein Anabolic Response to Increased Energy and Insulin Is Preserved in Poorly Controlled Type 2 Diabetes<sup>1,2</sup>

Jill A. Bell<sup>\*</sup>, Elena Volpi<sup>†</sup>, Satoshi Fujita<sup>†</sup>, Jerson G. Cadenas<sup>†</sup>, Melinda Sheffield-Moore<sup>‡</sup>, and Blake B. Rasmussen<sup>\*, \*\*, 3</sup>

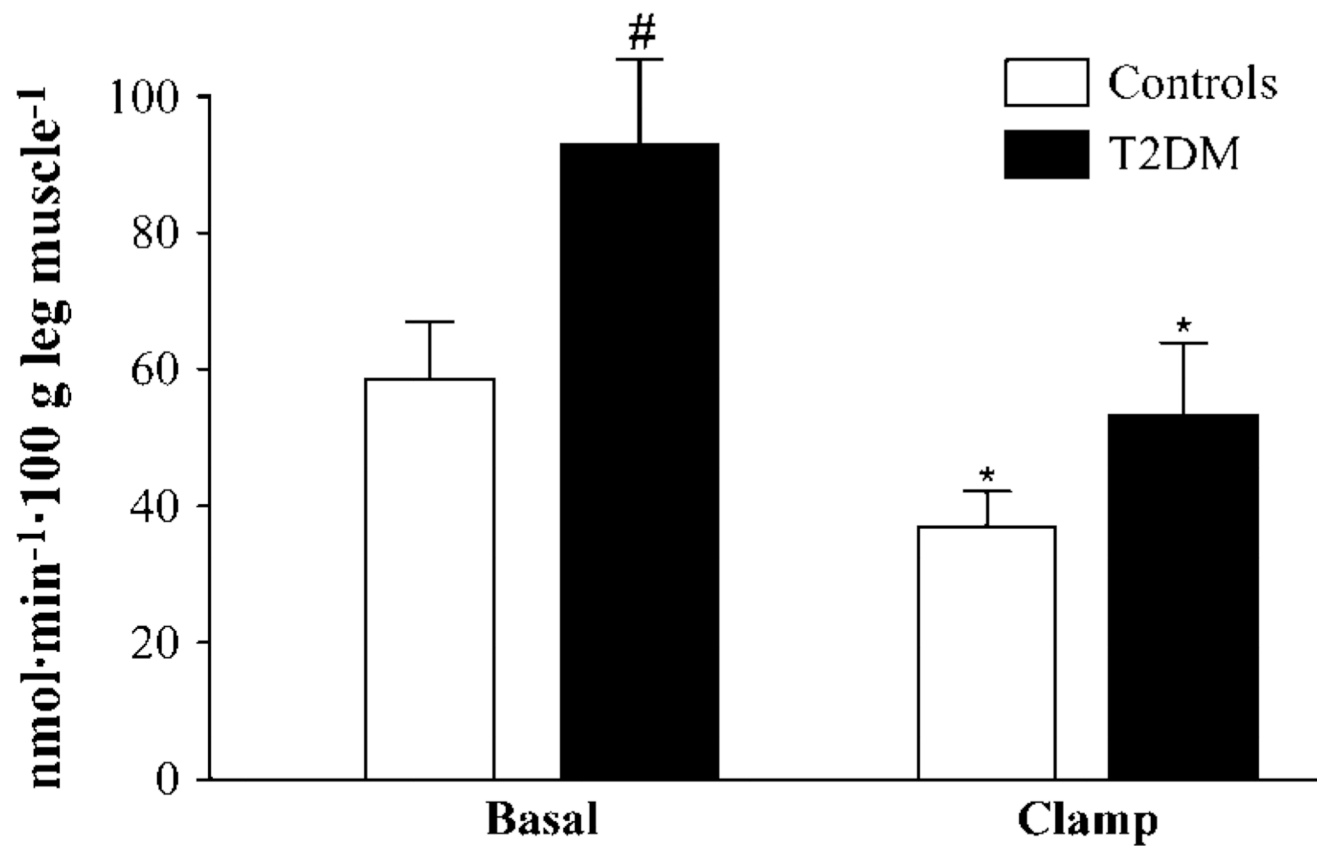


**FIGURE 1.**

Study design consisting of a basal postabsorptive period and a high energy-hyperinsulinemic clamp period. ICG, indocyanine green.

## Skeletal Muscle Protein Anabolic Response to Increased Energy and Insulin Is Preserved in Poorly Controlled Type 2 Diabetes<sup>1,2</sup>

Jill A. Bell<sup>\*</sup>, Elena Volpi<sup>†</sup>, Satoshi Fujita<sup>†</sup>, Jerson G. Cadenas<sup>†</sup>, Melinda Sheffield-Moore<sup>‡</sup>, and Blake B. Rasmussen<sup>\*,\*\*,3</sup>



## Longitudinal association of type 2 diabetes and insulin therapy with muscle parameters in the KORA-Age study

Uta Ferrari<sup>1</sup> · Cornelia Then<sup>1</sup> · Marietta Rottenkolber<sup>1</sup> · Canan Selte<sup>1</sup> · Jochen Seissler<sup>1</sup> · Romy Conzade<sup>2</sup> · Birgit Linkohr<sup>2</sup> · Annette Peters<sup>2,3</sup> · Michael Drey<sup>1</sup> · Barbara Thorand<sup>2,3</sup>

**Table 1** Baseline characteristics of the study participants stratified by diabetes status

	Total <sup>a</sup> (n = 731)		Diabetes <sup>a</sup> (n = 118 (16.1%))		No diabetes <sup>a</sup> (n = 613 (83.9%))		p value <sup>b</sup>
	Women	Men	Women	Men	Women	Men	
n (%)	360 (49.3)	371 (50.8)	58 (16.1)	60 (16.2)	311 (83.8)	302 (83.9)	0.9820 <sup>x</sup>
Age (years)	74.5 ± 6.2	74.7 ± 6.2	76.4 ± 5.8	75.4 ± 6.3	74.1 ± 6.3	74.5 ± 6.1	0.006/0.255
BMI (kg/m <sup>2</sup> )	28.3 ± 4.5	28.4 ± 3.9	31.0 ± 4.0	29.9 ± 4.4	27.7 ± 4.4	28.1 ± 3.7	<0.001/0.007
GS (kg)	20.7 ± 5.3	34.3 ± 7.6	20.2 ± 5.9	34.4 ± 7.5	20.7 ± 5.1	34.3 ± 7.7	0.409/0.602
SMI (kg/m <sup>2</sup> )	7.4 ± 0.9	10.1 ± 1.0	8.0 ± 1.0	10.5 ± 1.0	7.3 ± 0.9	10.0 ± 1.0	<0.001/<0.001
TUG (s)	10.5 ± 3.2	10.3 ± 2.9	11.8 ± 3.3	10.9 ± 2.6	10.3 ± 3.1	10.2 ± 3.0	<0.001/0.018
PASE: Total Score	119.1 ± 48.9	131.1 ± 59.7	112.7 ± 53.1	110.4 ± 57.6	120.3 ± 48.1	135.1 ± 59.4	0.173/0.002
HbA1c (%) (mmol/mol)	5.7 ± 0.5 (39)	5.7 ± 0.6 (39)	6.5 ± 0.7 (48)	6.5 ± 0.6 (48)	5.6 ± 0.3 (38)	5.6 ± 0.5 (38)	<0.001/<0.001
Insulin therapy only			4 (6.9%)	5 (8.3%)			
Oral anti-diabetics only			33 (56.9%)	38 (63.3%)			
Combination therapy <sup>y</sup>			7 (12.1%)	4 (6.7%)			
No anti-diabetic treatment			14 (24.1%)	13 (21.7%)			
Current smoker n (%)	16 (4.4)	21 (5.7)	2 (3.5)	4 (6.7)	14 (4.6)	17 (5.5)	0.624/0.925
Ex-Smoker n (%)	71 (19.7)	197 (53.1)	14 (24.1)	32 (53.3)	57 (18.9)	165 (54.6)	
Non-Smoker n (%)	273 (75.8)	153 (41.2)	42 (72.4)	24 (40.0)	231 (76.5)	129 (42.7)	
No chronic disease n (%)	33 (9.2)	54 (14.6)	3 (5.2)	5 (8.3)	30 (10.0)	49 (15.8)	0.017/0.006
1 chronic disease n (%)	107 (29.9)	115 (31.0)	10 (17.2)	11 (18.3)	97 (32.3)	104 (33.4)	
≥ 2 chronic diseases n (%)	218 (60.9)	202 (54.5)	45 (77.6)	44 (73.3)	173 (57.7)	158 (50.8)	

## Longitudinal association of type 2 diabetes and insulin therapy with muscle parameters in the KORA-Age study

Uta Ferrari<sup>1</sup> · Cornelia Then<sup>1</sup> · Marietta Rottenkolber<sup>1</sup> · Canan Selte<sup>1</sup> · Jochen Seissler<sup>1</sup> · Romy Conzade<sup>2</sup> · Birgit Linkohr<sup>2</sup> · Annette Peters<sup>2,3</sup> · Michael Drey<sup>1</sup> · Barbara Thorand<sup>2,3</sup>

		Model 1		Model 2	
		Regression coefficient (95% CI)	<i>p</i> value	Regression coefficient (95% CI)	<i>p</i> value
Δ GS	Total <i>n</i> = 90	− 1.2 (− 3.6 to 1.1)	0.294	− 1.6 (− 4.1 to 0.8)	0.190
	Men <i>n</i> = 47	− 3.0 (− 6.3 to 0.4)	0.081	− 3.4 (− 6.7 to 0.02)	0.051
	Women <i>n</i> = 43	0.1 (− 3.1 to 3.3)	0.927	0.4 (− 3.3 to 4.1)	0.830
Δ SMI	Total <i>n</i> = 88	0.5 (0.2–0.9)	<b>0.005</b>	<b>0.6 (0.3–0.9)</b>	<b>0.001</b>
	Men <i>n</i> = 45	0.5 (0.02–1.0)	<b>0.041</b>	0.5 (0.1–0.9)	<b>0.021</b>
	Women <i>n</i> = 43	0.5 (− 0.02 to 1.1)	0.060	0.8 (0.2–1.4)	<b>0.009</b>
Δ TUG	Total <i>n</i> = 79	1.4 (− 0.3 to 3.0)	0.096	1.6 (− 0.2 to 3.4)	0.086
	Men <i>n</i> = 43	0.1 (− 2.0 to 2.2)	0.946	0.4 (− 2.0 to 2.7)	0.758
	Women <i>n</i> = 36	2.7 (0.1–5.4)	<b>0.041</b>	4.3 (1.1–7.5)	<b>0.010</b>





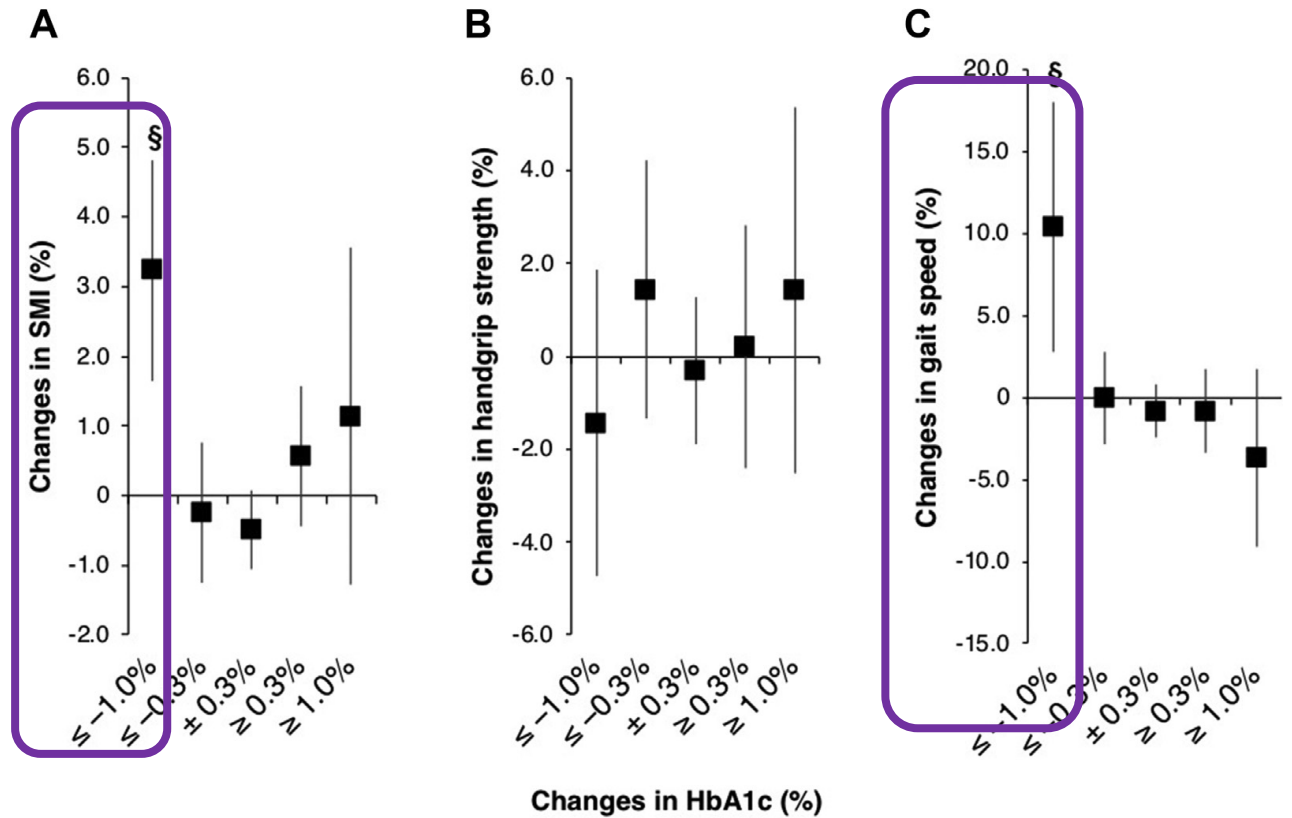
Original Study

Glycemic Control and Insulin Improve Muscle Mass and Gait Speed in Type 2 Diabetes: The MUSCLES-DM Study



Clinical Characteristics of Study Patients (n = 588)

	Baseline	1-y Follow-up	P Value
Age (y)	70.0 ± 8.9	71.0 ± 8.9	
Sex, male (%)		58.8	
Body weight (kg)	63.4 ± 12.0	63.5 ± 12.0	.84
BMI	24.7 ± 4.0	24.8 ± 3.9	.002
Waist circumference (cm)	90.5 ± 10.2	91.2 ± 10.1	<.001
Creatinine (mg/dL)	0.8 ± 0.3	0.9 ± 0.3	.003
HbA <sub>1c</sub> (%)	7.3 ± 1.2	7.1 ± 1.0	.001
HbA <sub>1c</sub> (mmol/mol)	55.8 ± 12.7	54.5 ± 10.5	.001
Regular exercise habit (%)	55.6	56.5	.68
Antihyperglycemic treatment			
Sulfonylureas (%)	28.1	27.4	.49
Glinides (%)	9.5	10.4	.32
Biguanides (%)	39.1	41.8	.014
Thiazolidinediones (%)	15.3	16.3	.32
DPP-4 inhibitors (%)	61.6	62.4	.52
SGLT-2 inhibitors (%)	12.8	15.3	.019
α-Glucosidase inhibitor (%)	14.7	16.7	.028
Number of oral drugs	1.8 ± 1.3	1.9 ± 1.3	<.001
GLP-1 analogs (%)	4.6	4.8	.81
Insulin (%)	25.9	26.5	.47
Muscle mass and function			
SMI	7.5 ± 1.2	7.5 ± 1.2	.80
Handgrip strength (kg)	28.4 ± 9.3	28.3 ± 9.4	.40
Gait speed (m/s)	1.18 ± 0.25	1.17 ± 0.25	.11
Sarcopenia (%)	6.3	7.8	.12





Original Study

## Glycemic Control and Insulin Improve Muscle Mass and Gait Speed in Type 2 Diabetes: The MUSCLES-DM Study

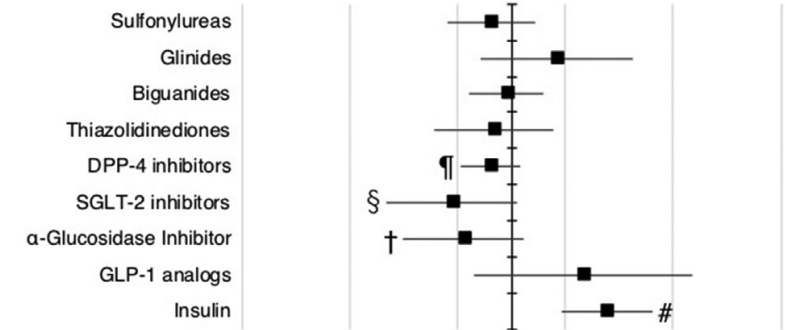


Linear Regression Analysis of Percentage Changes in SMI, Handgrip Strength, and Gait Speed During the Follow-up Period

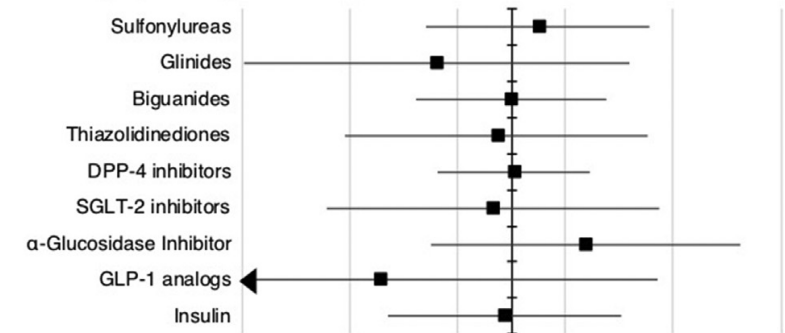
	Percentage Change During Follow-up Period					
	SMI		Handgrip Strength		Gait Speed	
	$\beta$	P Value	$\beta$	P Value	$\beta$	P Value
Age (years)	-0.021	.638	-0.178	<.001	-0.236	<.001
Sex (men)	0.183	.006	0.336	<.001	0.027	.490
BMI	0.173	.012	0.024	.574	-0.085	.033
Baseline SMI	-0.341	<.001				
Baseline handgrip strength (kg)			-0.439	<.001		
Baseline gait speed (m/s)					-0.421	<.001
HbA <sub>1c</sub> (%)	-0.003	.966	-0.096	.093	-0.067	.210
HbA <sub>1c</sub> ( $\geq 1\%$ decrease)	0.113	.027	-0.005	.914	0.145	.002
Regular exercise habit	0.015	.719	0.081	.054	0.050	.208
Antihyperglycemic drug use						
Insulin	0.115	.022				
DPP-4 inhibitors	-0.073	.104				
SGLT-2 inhibitors	-0.069	.125				
$\alpha$ -Glucosidase inhibitor	-0.068	.094				

n=558

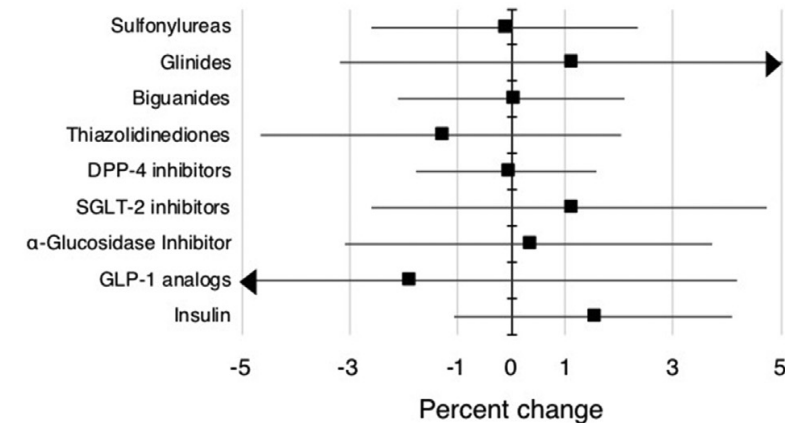
### A SMI



### B Handgrip strength



### C Gait speed





Original Study

Glycemic Control and Insulin Improve Muscle Mass and Gait Speed in Type 2 Diabetes: The MUSCLES-DM Study



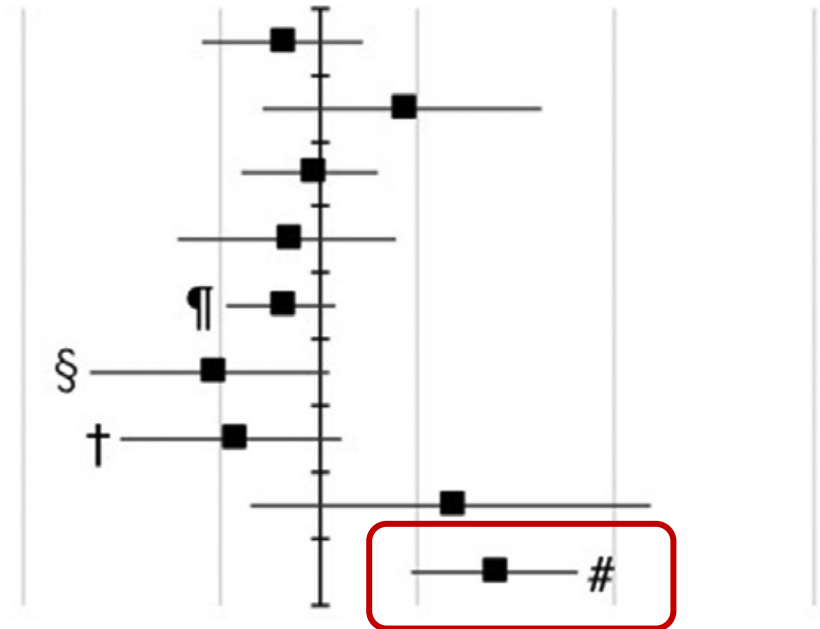
**SMI**

Linear Regression Analysis of Percentage Changes in SMI, Handgrip Strength, and Gait Speed During the Follow-up Period

	Percentage Change During Follow-up Period					
	SMI		Handgrip Strength		Gait Speed	
	$\beta$	P Value	$\beta$	P Value	$\beta$	P Value
Age (years)	-0.021	.638	-0.178	<.001	-0.236	<.001
Sex (men)	0.183	.006	0.336	<.001	0.027	.490
BMI	0.173	.012	0.024	.574	-0.085	.033
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$\alpha$ -Glucosidase inhibitor	-0.068	.094				

n=558

- Sulfonylureas
- Glinides
- Biguanides
- Thiazolidinediones
- DPP-4 inhibitors
- SGLT-2 inhibitors
- $\alpha$ -Glucosidase Inhibitor
- GLP-1 analogs
- Insulin

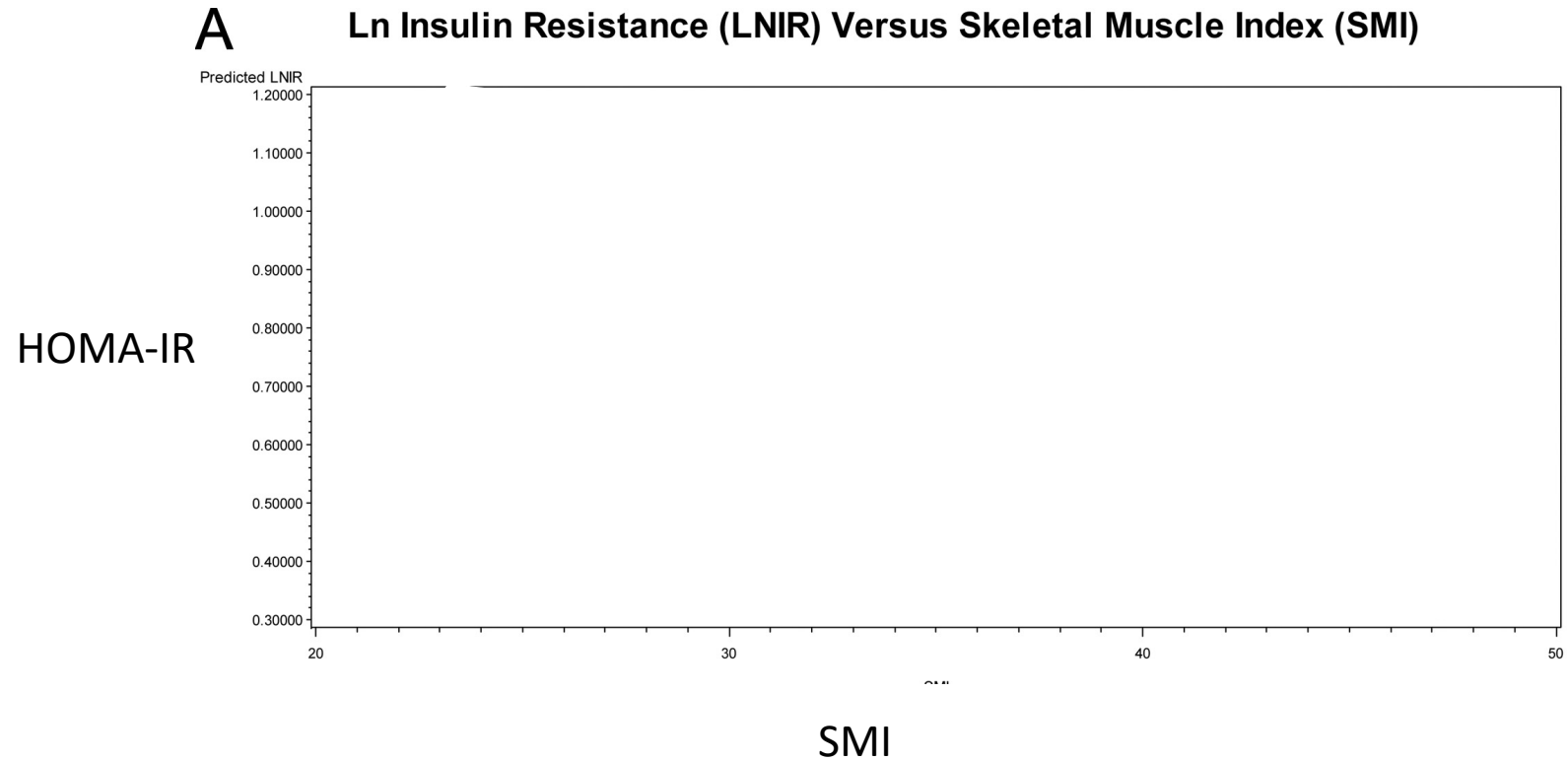


## Lower insulin level is associated with sarcopenia in community-dwelling frail and non-frail older adults

Yanxia Lu<sup>1</sup>, Wee Shiong Lim<sup>2,3</sup>, Xia Jin<sup>4</sup>,  
Ma Schwe Zin Nyunt<sup>5</sup>, Tamas Fulop<sup>6</sup>, Qi Gao<sup>5</sup>, Su Chi Lim<sup>7</sup>,  
Anis Larbi<sup>6,8</sup> and Tze Pin Ng<sup>5\*</sup>

		Sarcopenic subgroups			<i>P</i>
		Sarcopenia	Possible sarcopenia	Non-sarcopenia	
<b>Exploratory study</b>					
Fasting insulin (pg/ml)	453.57 ± 318.30	375.90 ± 233.30 <sup>*+</sup>	489.34 ± 328.09	565.32 ± 453.61	0.023
C-peptide (ng/ml)	1.54 ± 0.85	1.35 ± 0.63	1.64 ± 0.81	1.72 ± 1.39	0.067
Fasting glucose (mmol/L)	5.33 ± 1.28	5.35 ± 1.37	5.36 ± 1.29	5.16 ± 0.80	0.755
IGF-1 (ng/ml)	1.02 ± 0.74	1.02 ± 0.72	0.99 ± 0.76	1.12 ± 0.77	0.751
Leptin (ng/ml)	11.07 ± 10.05	8.92 ± 9.49 <sup>+</sup>	13.16 ± 10.72	9.70 ± 7.27	0.029
Active ghrelin (pg/ml)	3.90 ± 4.51	3.47 ± 3.63	3.89 ± 4.72	5.38 ± 6.00	0.255
<b>Validation study</b>					
Fasting insulin (pg/ml)	307.92 ± 251.07	232.87 ± 206.93 <sup>*+</sup>	395.47 ± 251.30	351.33 ± 303.28	< 0.001
C-peptide (ng/ml)	1.03 ± 0.45	0.90 ± 0.36 <sup>*+++</sup>	1.17 ± 0.49	1.14 ± 0.53	< 0.001
Fasting glucose (mmol/L)	6.19 ± 1.57	6.23 ± 1.70	6.27 ± 1.53	5.94 ± 1.18	0.596
IGF-1 (ng/ml)	14.60 ± 7.15	15.36 ± 7.59	13.75 ± 6.97	14.47 ± 6.46	0.514
Leptin (ng/ml)	15.53 ± 16.23	10.96 ± 9.27 <sup>+++</sup>	23.27 ± 21.87 <sup>**</sup>	13.34 ± 13.39	< 0.001
Active ghrelin (pg/ml)	21.57 ± 16.96	22.39 ± 16.95	21.97 ± 16.70	18.42 ± 17.67	0.510

\*\* *P* < 0.01, \* *P* < 0.05 vs. the non-sarcopenia group; +++ *P* < 0.001, + *P* < 0.05 vs. the possible sarcopenia group.

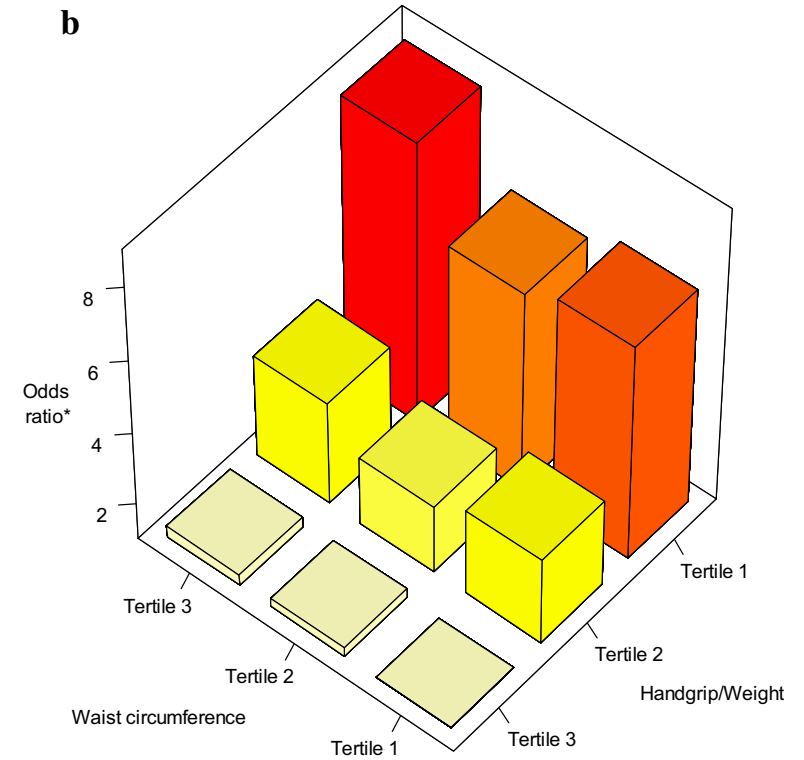
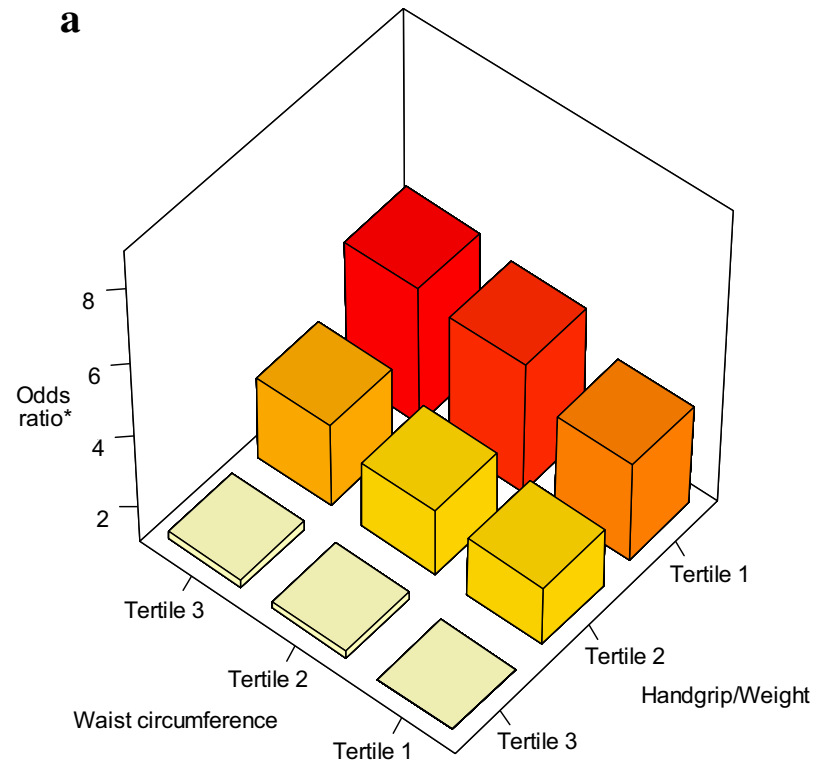


Each 10% increase in SMI was associated with 11% relative reduction in HOMA-IR  
(95% confidence interval, 6 –15%)



ORIGINAL INVESTIGATION Open Access

The prediction of Metabolic Syndrome alterations is improved by combining waist circumference and handgrip strength measurements compared to either alone

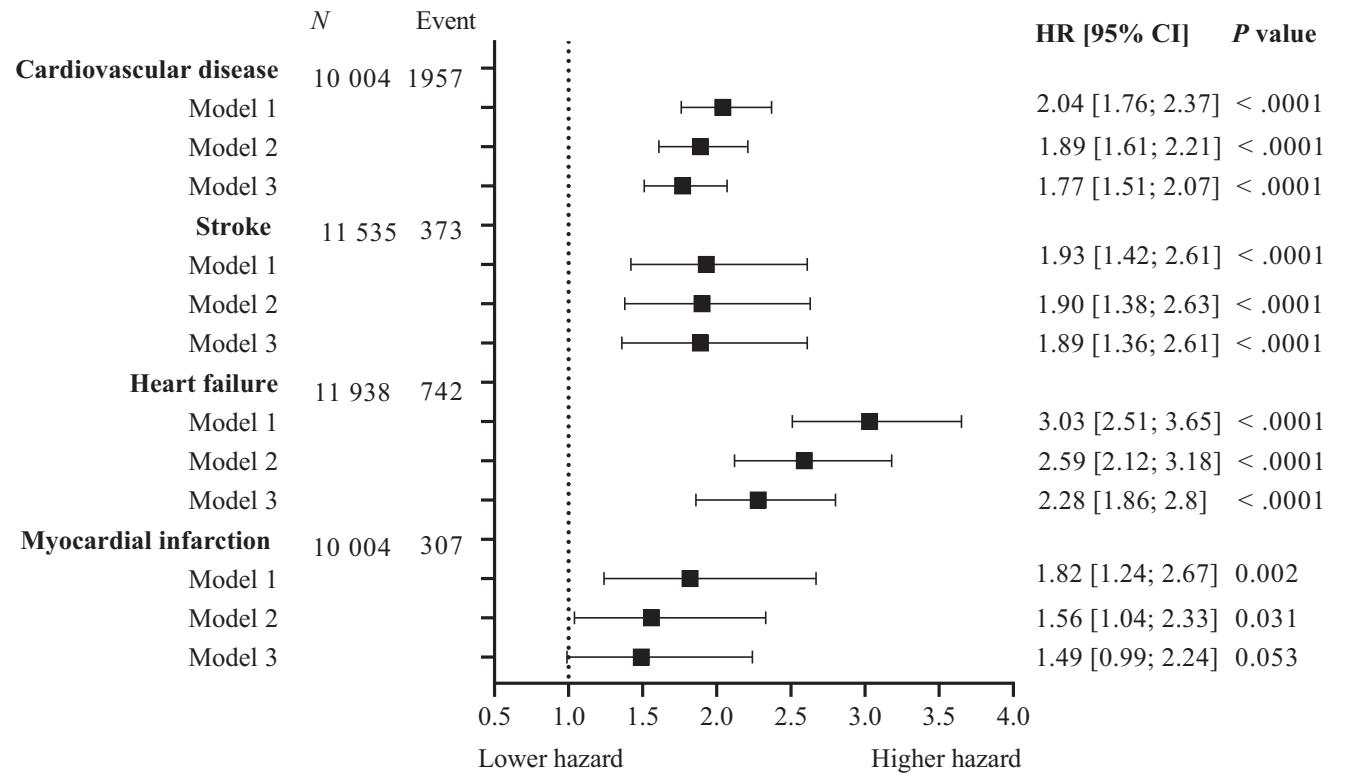


Received: 20 July 2023 | Revised: 22 September 2023 | Accepted: 4 October 2023  
 DOI: 10.1111/dom.15338

ORIGINAL ARTICLE

WILEY

**In people with type 2 diabetes, sarcopenia is associated with the incidence of cardiovascular disease: A prospective cohort study from the UK Biobank**







**TABLE 2** The RAP among sarcopenia compared with non-sarcopenia.

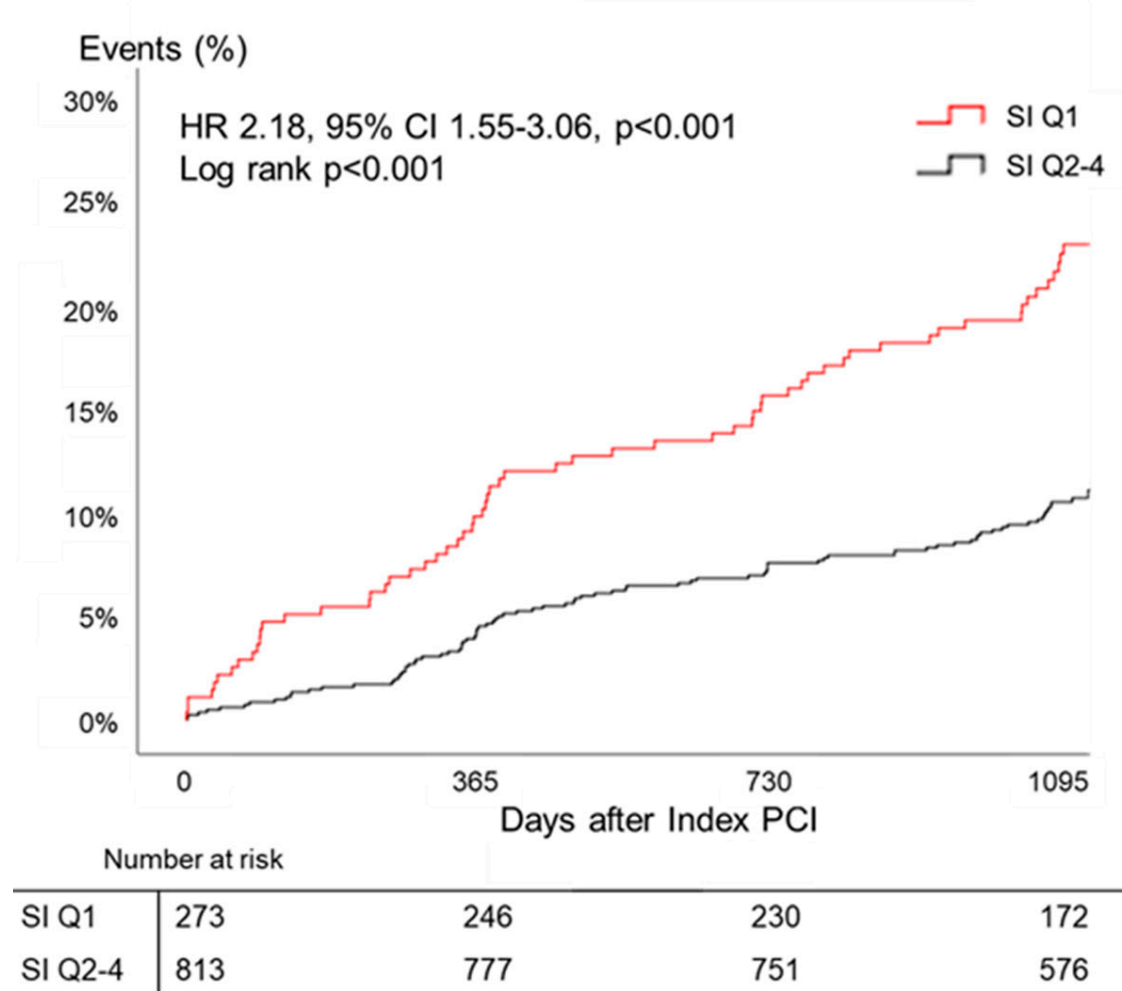
Outcomes	RAP (95% CI)
Cardiovascular disease	14.5 (13.1; 15.6)
Stroke	13.7 (10.8; 15.1)
Heart failure	13.7 (13.3; 14.0)
Myocardial infarction	12.8 (2.4; 16.3)

Article

## Sarcopenia Index as a Predictor of Clinical Outcomes in Older Patients with Coronary Artery Disease

Hak Seung Lee , Kyung Woo Park <sup>\*</sup>, Jeehoon Kang , You-Jeong Ki , Mineok Chang, Jung-Kyu Han, Han-Mo Yang , Hyun-Jae Kang, Bon-Kwon Koo and Hyo-Soo Kim

### A. MACE
















Volumen 12, número 4 de 2025  
<https://doi.org/10.53853/encr.12.4.935>

REVISTA  
COLOMBIANA de Endocrinología  
Diabetes & Metabolismo

## Consensos

### Consenso colombiano basado en evidencia y en la opinión de expertos en el manejo integral de pacientes con diabetes *mellitus* tipo 2 y sarcopenia

Jorge Castillo  <sup>1</sup>, Martín Vásquez <sup>2</sup>, Jhon Jairo Duque <sup>3</sup>, Diana Carolina Díaz Tribaldos <sup>4</sup>,  
Juan Carlos Galvis <sup>5</sup>, Jaime Ibarra <sup>6</sup>, Edgar Castro <sup>7</sup>, Carlos Rosselli <sup>8</sup>, Angélica Veloza <sup>9</sup>,  
Karen Cárdenas–Garzón <sup>10</sup>, Juan Pablo Zuluaga Peña <sup>10</sup>, Julio Ricardo Zuluaga Peña <sup>10</sup>



@eldoctorcastillo

**Insulina:** efecto positivo. Se ha informado que previene la disminución de la masa muscular en las extremidades inferiores, no afecta la velocidad de la marcha y está asociada con una menor probabilidad de reducción del índice de masa muscular esquelética. Se recomienda dado su efecto sobre la ganancia de peso (26).

**Inhibidores de la DPP-4 (iDPP-4):** efecto positivo o neutral. La evidencia sugiere que el uso de estos medicamentos impacta positivamente en los parámetros de sarcopenia.

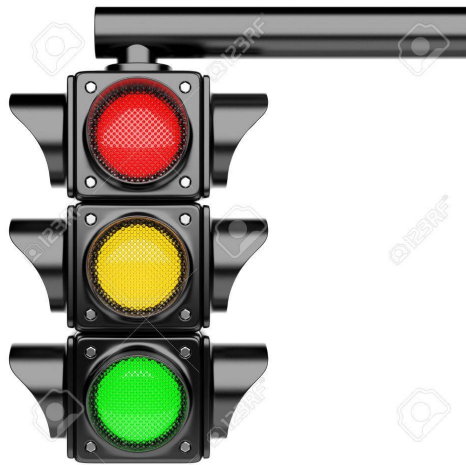
**Metformina:** presenta un efecto positivo o neutral. Su uso se asocia con una menor pérdida de masa muscular esquelética total y apendicular, así como con una menor disminución en la velocidad de la marcha habitual y no altera los resultados de la prueba de fuerza de agarre.

**Agonistas del receptor GLP-1 (aGLP-1):** efecto neutral. Los estudios de los GLP-1 en las dosis recomendadas para el manejo del paciente con diabetes no reportan cambios significativos en la masa muscular esquelética.

**Inhibidores del SGLT-2 (iSGLT-2):** efecto neutro o incierto. No se han reportaron cambios significativos en la masa muscular esquelética.

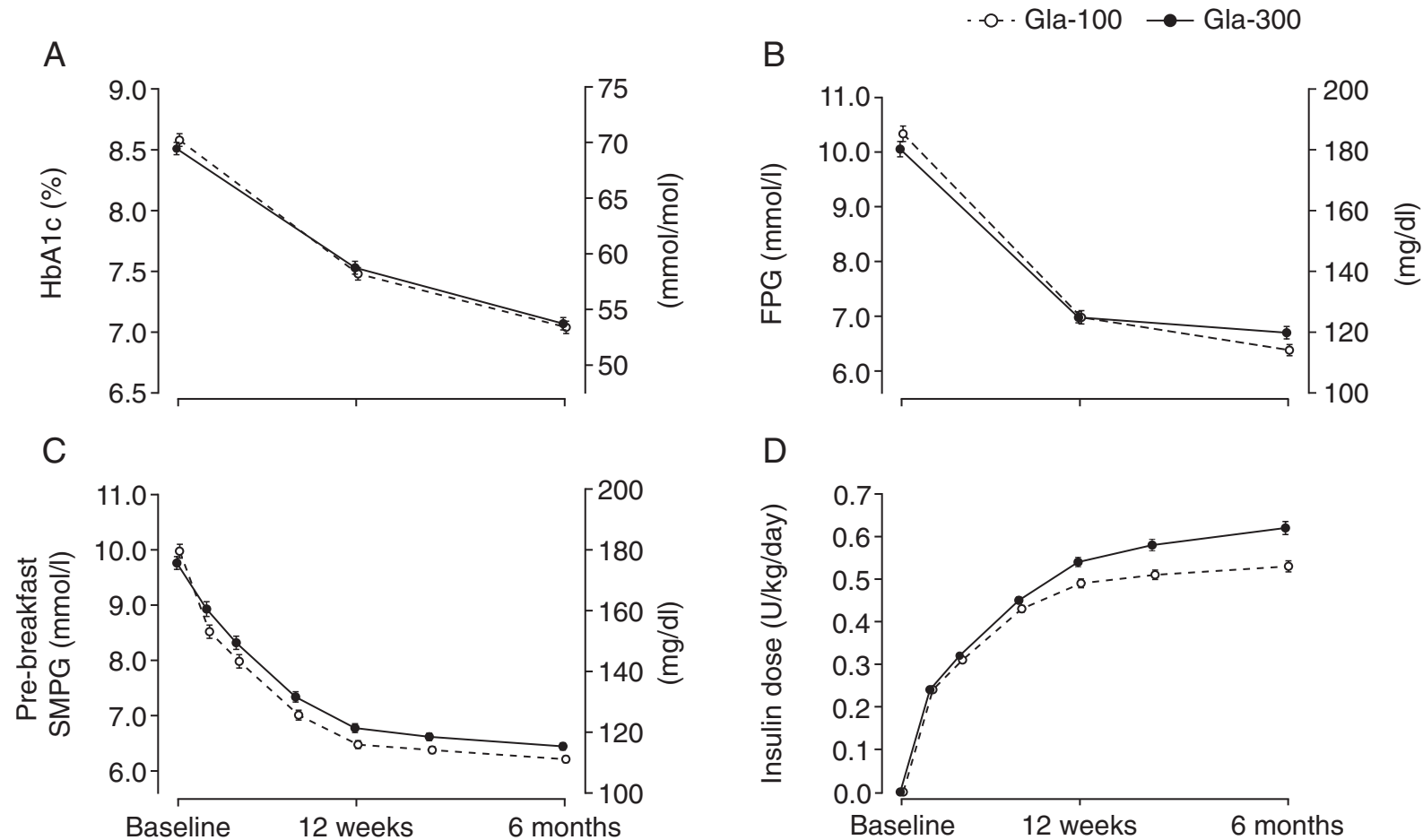
**Sulfonilureas y glinidas:** tienen un efecto negativo, aunque la evidencia es limitada. Se han asociado con una disminución en la masa muscular esquelética, el índice de músculo esquelético, la fuerza muscular y la velocidad de la marcha.

**Tiazolidinedionas:** efecto negativo. Están relacionadas con una reducción significativa de la masa muscular y de la velocidad de la marcha habitual.

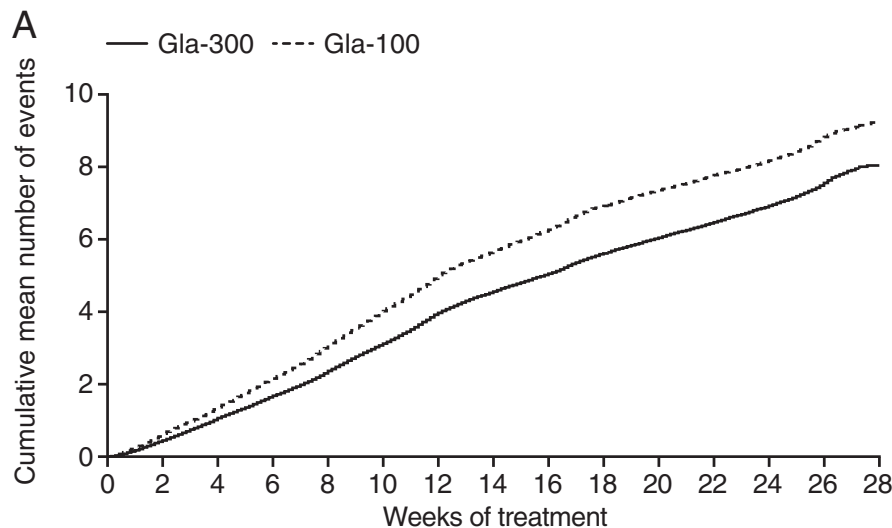


## New insulin glargine 300 U/ml compared with glargine 100 U/ml in insulin-naïve people with type 2 diabetes on oral glucose-lowering drugs: a randomized controlled trial (EDITION 3)

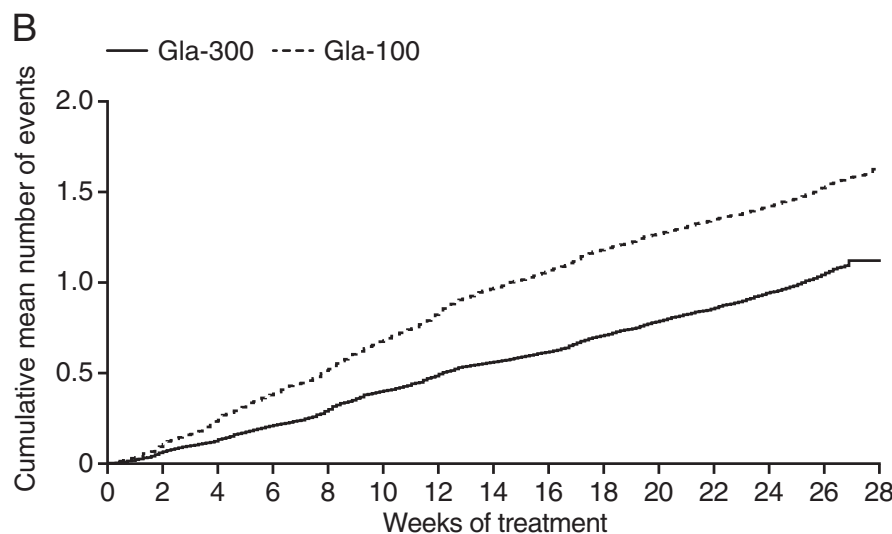
G. B. Bolli<sup>1</sup>, M. C. Riddle<sup>2</sup>, R. M. Bergenstal<sup>3</sup>, M. Ziemien<sup>4</sup>, K. Sestakauskas<sup>5</sup>, H. Goyeau<sup>6</sup>, P. D. Home<sup>7</sup> & on behalf of the EDITION 3 study investigators



Any time of day (24 h)



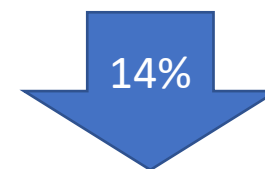
Nocturnal (00:00–05:59 h)



**Patient-level meta-analysis of the EDITION 1, 2 and 3 studies: glycaemic control and hypoglycaemia with new insulin glargine 300 U/ml versus glargine 100 U/ml in people with type 2 diabetes**

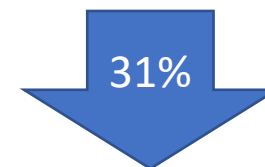
R. Ritzel<sup>1</sup>, R. Roussel<sup>2,3,4</sup>, G. B. Bolli<sup>5</sup>, L. Vinet<sup>6</sup>, C. Brulle-Wohlhueter<sup>7</sup>, S. Glezer<sup>7</sup> & H. Yki-Järvinen<sup>8</sup>

RR 0.86, 95% CI 0.77–0.97; p = 0.0116



Tasa de eventos anualizada < 70 mg/dl

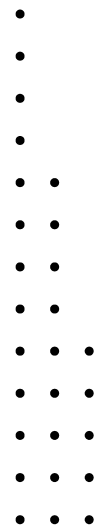
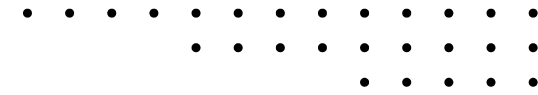
2.10 vs 3.06, RR 0.69, 95% CI 0.57–0.84; p = 0.0002



# Diabetes, sarcopenia u obesidad ¿cuál es el enemigo?



@eldoctorcastillo



# Gracias

 @ eldoctorcastillo

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