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Congreso Nacional de la Sociedad
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LA VISIÓN GLOBAL DE LA PERSONA ENFERMA



19-21 Noviembre 2014

Auditorio y Centro de Congresos Víctor Villegas
Murcia

Pacientes con enfermedades endocrinológicas distintas a la diabetes

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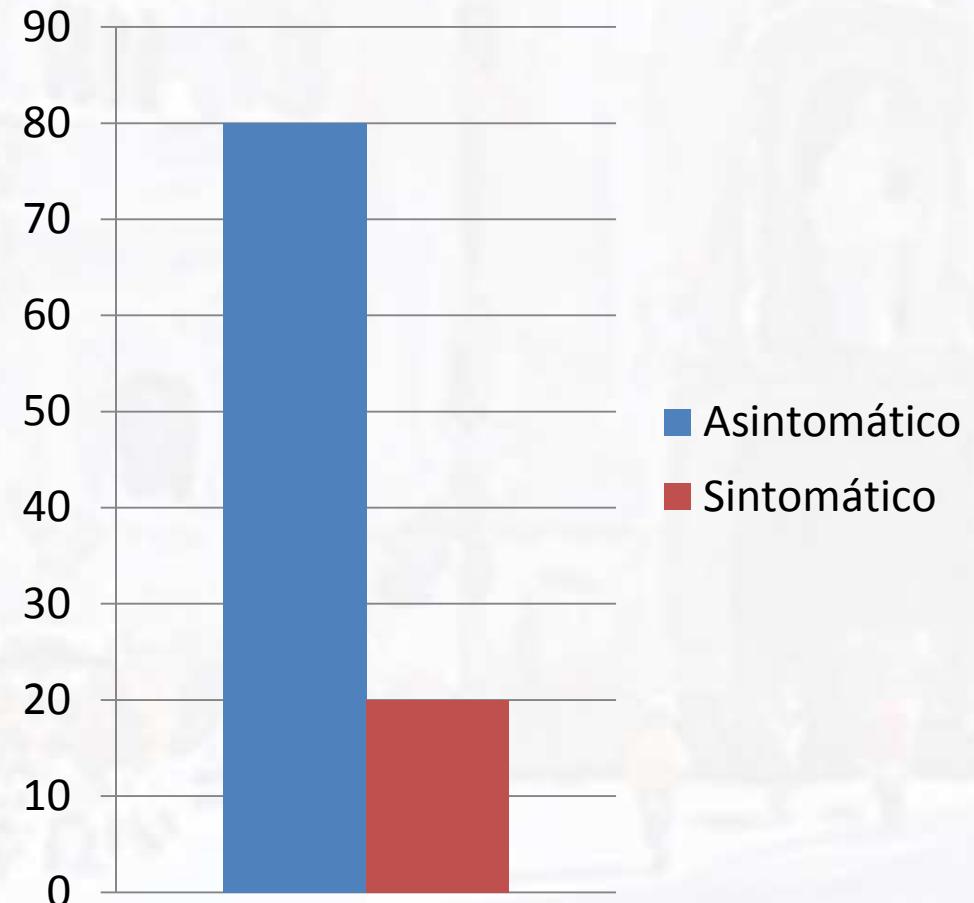
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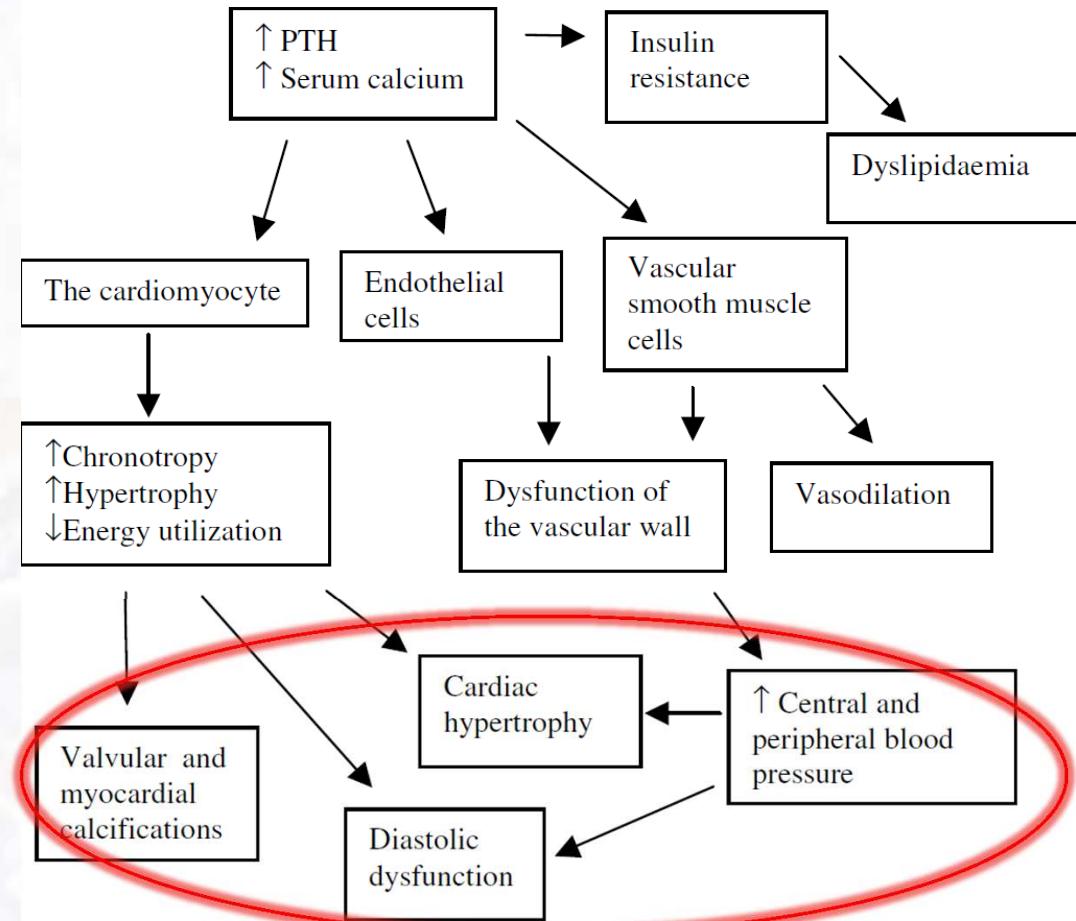
- **HIPERPARATIROIDISMO PRIMARIO**
- **SÍNDROME DE CUSHING**
- **HIPOTIROIDISMO SUBCLÍNICO**

Hiperparatiroidismo primario

- Hipercalcemia
- Nefrolitiasis
- Afectación ósea
- Hipertensión
- Afectación digestiva
- Manifestaciones neuropsiquiátricas



Hiperparatiroidismo primario



European Heart Journal (2004) 25, 1776–1787



Hiperparatiroidismo primario

Table 1 Clinical data regarding mortality studies among pHPT patients

Author (ref no.)	N (controls)	N (patients)	Mean age (patients)	Serum-calcium (mmol/l)	Mean follow-up time (years)	Relative risk of death vs. controls	Risk ratio for CVD
Hedbäck ⁵	ES	896	57 ± 13	3.03 ± 0.50/ 2.81 ± 0.27 ^c	12.9 ± 6.1	1.67 ($p < 0.001$)	1.66 ($p < 0.001$)
Hedbäck ⁶	ES	915	61 ± 14	NP	3.5 (0–8)	1.30 ($p = 0.0099$)	1.71 95% CI (1.34–2.15) ^d
		3446	65 ± 13	NP	3.6 (0–8)	1.61 ($p < 0.001$)	1.85 95% CI (1.62–2.11) ^d
Palmer ²⁶	ES	441	58 ± 13	2.87 ± 0.30	7.7	1.06	1.17 ($p = 0.06$) ^a
Uden ²⁹	NP	282	59 (16–88)	2.91 (2.44–5.31)	8 ± 2	NP	1.18 (NS)
Palmer ³¹	344	172	59 ± 12	2.72 ± 0.14	14	2.21 ($p = 0.0135$)	1.43 (NS)
Wermers ³⁴	ES	435	57 (16–89)	2.72 ± 0.12	NP	0.69 95% CI (0.57–0.83) ^d	0.60 95% CI (0.45–0.79) ^{d,b}



Hiperparatiroidismo primario

Table 2 Clinical data regarding LVM studies

Author (ref no.)	N (patients)	N (controls)	Mean age of the patients (years)	BP patients vs controls	Calcium (mmol/l)	PTH (pg/ml)	LVMI patients	LVMI controls	P LVMI
Längle ⁷⁰			57 ± 16						
Asymptomatic	7	NP		NP	2.83 ± 0.17	123 ± 47	LVMI NP	LVMI NP	
Minimal symptoms	48	NP		NP	3.03 ± 0.36	174 ± 127	LVMI NP	LVMI NP	
Symptomatic	77	NP		NP	3.08 ± 0.47	279 ± 263	LVMI NP	LVMI NP	
Stefenelli ¹⁹	69	NP	64 ± 10	NP	2.98 ± 0.41	209 ± 120	11.5 ± 2.0 ^c	10.1 ± 2.2 ^{ce}	<0.05
Stefenelli ²⁰	54	50	61 ± 11	NS	2.99 ± 0.28	231 ± 208	12.2 ± 1.4 ^{cd}	10.8 ± 1.7 ^{cd}	<0.001
Dalberg ⁷¹	44	23	61 (52–70) ^a	NS	2.78 (2.69–2.92) ^a	82 (73–102) ^a	11.0 ^c	9.5 ^c	0.03
Piovesan ²¹	43	43	60 ± 13	NS	2.81 ± 0.30	161 ± 73	136.1 ± 24	114 ± 15	<0.05
Almqvist ²²	25	25	70 ± 8/68 ± 9 ^b	NS	2.62 ± 0.11	82 ± 34	112 ± 27 ^b	131 ± 34 ^b	<0.05
Dominiczak ¹⁷	23	23	54 ± 3	NS	2.73	5.8 (units/l)	123 ± 10	100 ± 6	0.03
Nilsson ⁷⁵	30	30	64 ± 12	NS	2.64 ± 0.11	77 ± 24	124 ± 5	114 ± 4	0.06
Barletta ⁵²	14	20	60 ± 11	NS	2.88 ± 0.26	215 ± 188	91 ± 18	89 ± 19	NS
Nuzzo ⁹	20	20	53 ± 9	NS	3.04 ± 0.22	280 ± 165	113 ± 13	111 ± 11	NS



Hiperparatiroidismo primario

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Table 3 Clinical data regarding diastolic function among pHPT patients

Author (ref no.)	n	BP pre-op vs. post-op	E/A pre-op	E/A post-op	p
Dalberg ⁷¹	44	$p < 0.01^c$	a	a	<0.02
Näppi ²³	15	NP	1.028 ± 0.373	0.909 ± 0.278	NS
Stefenelli ¹⁹	16	NS	NP	NP	NS
I-L Nilsson ⁷⁵	30	NS	0.99 ± 0.07	1.0 ± 0.06	NS
Ohara ⁷⁸	14	NS	1.51 ± 0.51^b	1.12 ± 0.32^b	<0.01
Almqvist ²²	50	NS	0.81 ± 0.20	0.95 ± 0.38	<0.05
Barletta ⁵²	14	NS	0.94 ± 0.18	0.96 ± 0.16	NS



Hiperparatiroidismo primario

Table 4 Clinical data regarding cardiac calcifications among pHPT patients

Author (ref no.)	<i>n</i>	Aortic valve (%)	Mitral valve (%)	Myocardial (%)	Calcium (mmol/l)	PTH (pg/ml)
Längle ⁷⁰	132					
Asymptomatic	7	14	14	86	2.83 ± 0.17	123 ± 47
Minimal symptoms	48	33	4	58	3.03 ± 0.36	174 ± 127
Symptomatic	77	14	16	61	3.08 ± 0.47	279 ± 263
Stefenelli ²⁰	54	63	49	69	2.99 ± 0.28	231 ± 208
Stefenelli ¹⁹	69	54	39	74	2.98 ± 0.41	209 ± 119



Hiperparatiroidismo primario

Table I. Symptoms and end-organ damage in PHPT patients ($n = 56$)

Symptoms	%
Bone pain	42
Bone swelling/deformity	11
Fractures	10
Osteoporosis (t score < -2.5)	33
Muscle weakness	23
Nephrocalcinosis/Nephrolithiasis	25
Behavioral problems	14
Acute or chronic pancreatitis	9
Acid peptic disease	28

	Controls (n = 25)	PHPT (n = 56)	P value
s. calcium, mg/dL	8.7 ± 0.5	11.5 ± 1.3	.001
s. NT-proBNP, pg/mL	58 ± 49	$463 \pm 1,130$.002
Two-dimensional echocardiographic indices			
LVEDD, mm	43.7 ± 5.2	43.5 ± 6.5	.95
LVESD, mm	26.6 ± 4.3	26.6 ± 6.9	.63
LVEDV, mL	63.6 ± 18.3	67.3 ± 23.0	.63
LVESV, mL	27.1 ± 8.5	25.2 ± 11.3	.34
LVM, g	148.9 ± 44.1	192 ± 70.1	.006
IVS, mm	9.0 ± 1.6	10.8 ± 2.5	.001
PW, mm	8.6 ± 2.2	9.9 ± 2.0	.004
LVEF, %	64 ± 6	62 ± 9	.10
E/A ratio	1.3 ± 0.4	1.1 ± 0.5	.01
IVRT, ms	89.5 ± 25	87 ± 23	.44
Calcification, %	8	26.7	.05
Tissue Doppler imaging, m/s			
E α	0.13 ± 0.05	0.11 ± 0.0	.02
A α	0.11 ± 0.04	0.26 ± 0.9	.19
S	0.74 ± 3.1	0.14 ± 0.1	.33
Vasodilatory response studies, %			
FMD	12 ± 6	9 ± 9	.03
NMD	24 ± 11	20 ± 18	.01

(Surgery 2013;154:1394-404.)

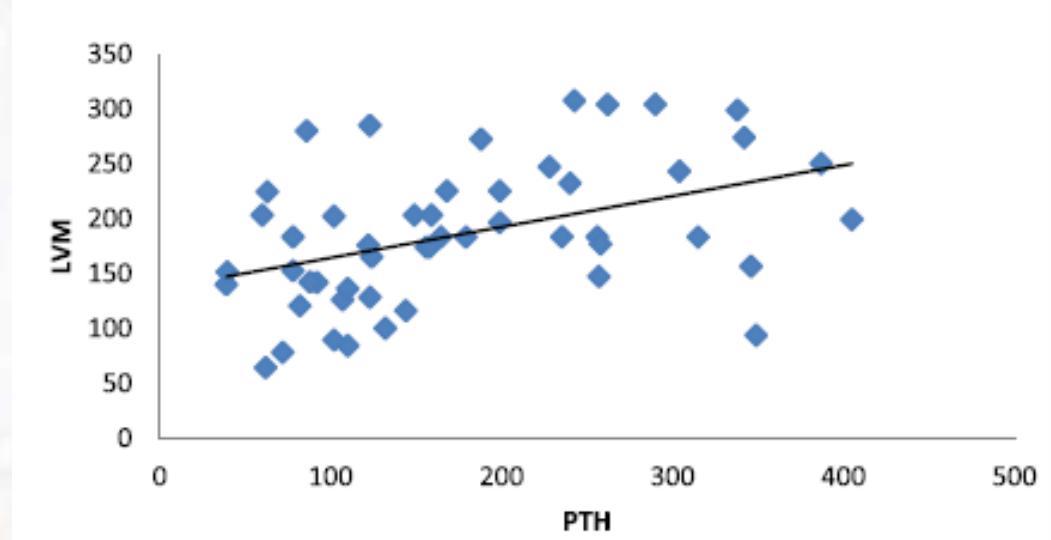


Hiperparatiroidismo primario



Table IV. Pearson's product moment correlation analysis on biochemical parameters and left ventricular mass in PHPT patients

	<i>Left ventricular mass</i>	
	<i>R value</i>	<i>P value</i>
s. PTH	0.502	<.001
s. Calcium (total)	0.061	.680
s. 25 hydroxy vitamin D	-0.194	.177
s. NT-pro BNP	0.039	.790



(Surgery 2013;154:1394-404.)



Hiperparatiroidismo primario

Table VII. Linear changes in biochemical, two-dimensional echocardiographic, tissue Doppler imaging, and vasodilatory response indices in hypertensive primary hyperparathyroidism patients after parathyroidectomy

	Baseline value	3 month after PTx	P value	6 month after PTx	P value
s. calcium, mg/dL	11.5 ± 1.2	9.1 ± 1.1	<.001	9.1 ± 0.7	<.001
s. NT-proBNP, pg/mL	399 ± 695	236 ± 363	.211	231 ± 372	.272
Two-dimensional echocardiographic indices					
LVEDD, mm	45.2 ± 4.5	43.6 ± 7.3	.020	42.1 ± 3.9	.001
LVEDS, mm	27.2 ± 3.7	27.7 ± 3.9	.605	27.0 ± 4.1	.872
LVEDV, mL	67.5 ± 25.4	65.3 ± 22.6	.551	68.9 ± 25.6	.626
LVESV, mL	25.3 ± 11.5	25.5 ± 15.4	.944	21.6 ± 11.4	.037
IVS, mm	11.8 ± 2.3	11.6 ± 2.0	.454	11.3 ± 2.9	.189
PW, mm	10.2 ± 1.6	10.3 ± 1.7	.806	9.7 ± 1.6	.150
LVEF, %	60 ± 9	65 ± 12	.117	68 ± 14	.011
Stroke volume, mL	42.0 ± 17.1	38.7 ± 18.9	.535	47.3 ± 19.6	.071
LVM, g	215.0 ± 59.9	202.7 ± 61.3	.059	171.3 ± 63.6	.014
E/A ratio	1.1 ± 0.4	1.1 ± 0.4	.043	1.2 ± 0.4	.001
IVRT, ms	88.3 ± 23.1	89.7 ± 18.2	.778	91.1 ± 15.9	.471
Tissue Doppler imaging, m/s					
Eα	0.098 ± 0.04	0.11 ± 0.04	.034	0.12 ± 0.02	.003
Aα	0.58 ± 1.7	0.10 ± 0.02	.31	0.10 ± 0.02	.02
S	0.17 ± 0.21	0.10 ± 0.03	.26	0.11 ± 0.04	.09
Vasodilatory response studies, &					
FMD	10 ± 10	12 ± 8	.28	11 ± 6	.77
NMD	21 ± 16.5	21 ± 16	.56	26 ± 17	.036

(Surgery 2013;154:1394-404.)



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Síndrome de Cushing

Clinical features of Cushing's syndrome

- Obesity
- Moon facies
- Hypertension
- 'Buffalo hump'
- Thin skin^a
- Hirsutism
- Oligomenorrhoea/amenorrhoea
- Purple striae
- Impaired glucose tolerance/diabetes mellitus
- Proximal muscle atrophy^a
- Psychiatric disturbance
- Osteoporosis
- Bruising^a
- Acne
- Hypokalaemia



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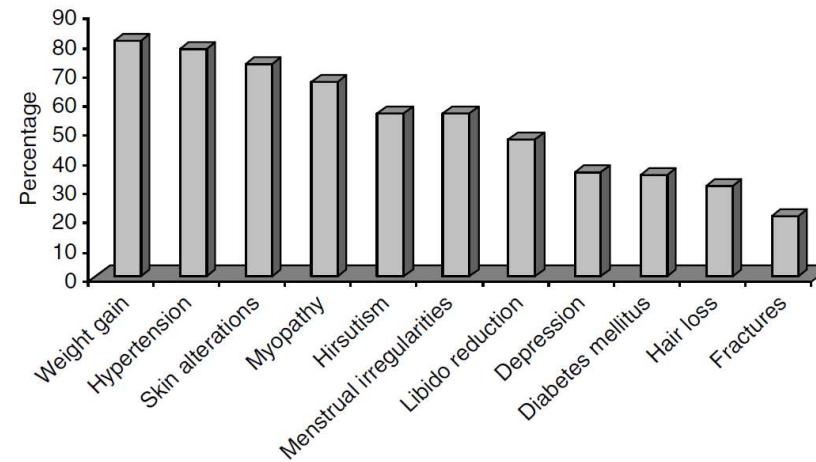
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MurciaEUROPEAN JOURNAL OF ENDOCRINOLOGY (2011) **165**

	PIT-CS	ADR-CS	ECT-CS	OTH-CS	Overall
General practitioner	198/260 (76)	90/114 (79)	18/22 (82)	5/10 (50)	311/406 (77)
Diabetologist	58/236 (25)	19/98 (19)	9/22 (41) [†]	2/9 (22)	88/365 (24)
Gynaecologist	47/193 (24)*	21/85 (25)*	1/11 (9)	2/8 (25)	71/297 (24)
Psychiatrist/psychologist	28/226 (12)	10/97 (10)	2/22 (9)	2/10 (20)	42/355 (12)
Rheumatologist/orthopaedist	25/224 (11)	10/96 (10)	2/22 (9)	2/10 (20)	39/352 (11)
Dermatologist	18/227 (8)	5/97 (5)	1/22 (5)	2/10 (20)	26/356 (7)
Other ^a	121/229 (53)	47/90 (52)	10/18 (56)	6/9 (67)	184/346 (53)

European Journal of Endocrinology 165 383-392

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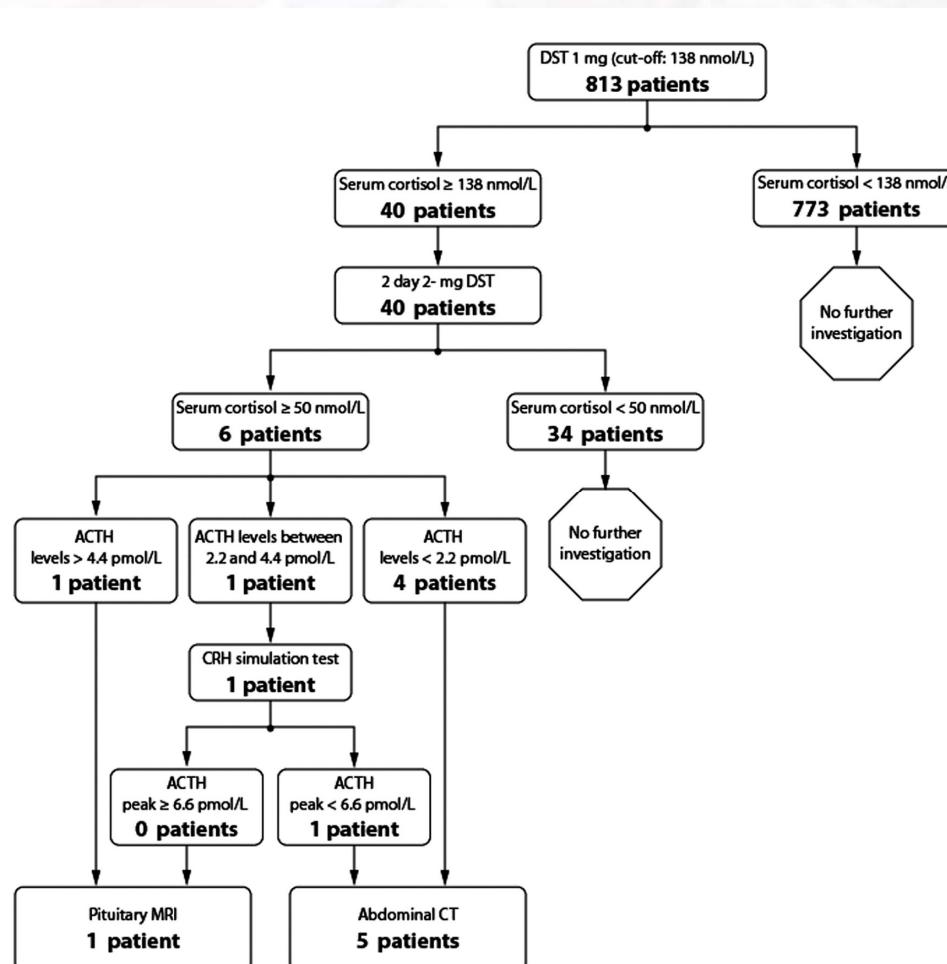
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TABLE 2. Comparison of patients with post-DST cortisol 5 µg/dl or less (DST suppressors) and post-DST cortisol greater than 5 µg/dl (DST nonsuppressors)

	DST suppressors (n = 773)	DST nonsuppressors (n = 34) ^a	P value
Age (yr)	58.6 ± 8.8	56.4 ± 9.8	NS
BMI (kg/m ²)	32.1 ± 6.1	30.9 ± 4.6	NS
Duration of disease (yr)	9.8 ± 7.9	8.9 ± 5.2	NS
Fasting glucose (mg/dl)	172.5 ± 65.1	236.8 ± 100.8	<0.0001
HbA1c (%)	8.4 ± 1.9	9.2 ± 2.0	0.01
Systolic blood pressure (mm Hg)	137.9 ± 17.2	147.7 ± 16.4	0.04
Diastolic blood pressure (mm Hg)	82.6 ± 10.1	80.7 ± 10.9	NS



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Incidence and Late Prognosis of Cushing's Syndrome: A Population-Based Study

TABLE 2. Standard mortality ratios (SMR) in patients with Cushing's syndrome

Diagnosis	No. of patients	Age ^a (yr), median, range	No. of deaths	Time at risk (yr)	Expected no. deaths	SMR (95% CI)	P ^b
Cushing's disease (etiology proven)	73	41.1, 7.6–69.7	7	560.0	4.12	1.70 (0.68–3.50)	0.14
Cushing's disease (etiology unproven)	26	51.1, 24.9–74.1	11	153.1	0.96	11.48 (5.73–20.5)	<0.001
Adrenal adenoma	37	38.3, 3.6–77.7	4	255.2	1.15	3.48 (0.95–8.90)	0.04

CI, Confidence interval.

^a Age at first admission.

^b Compared to the expected from national mortality rates.

J Clin Endocrinol Metab 86: 117-123,



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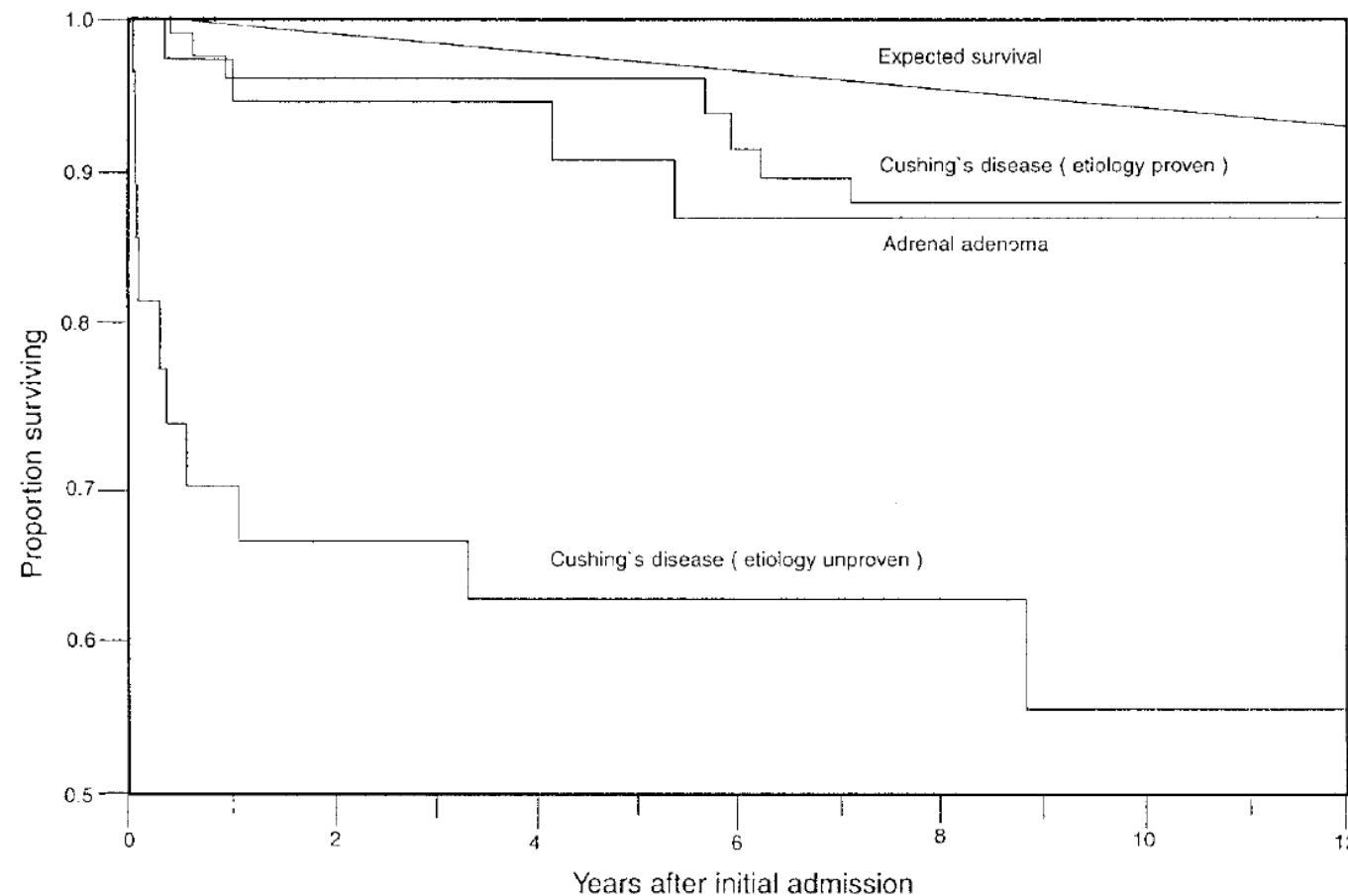
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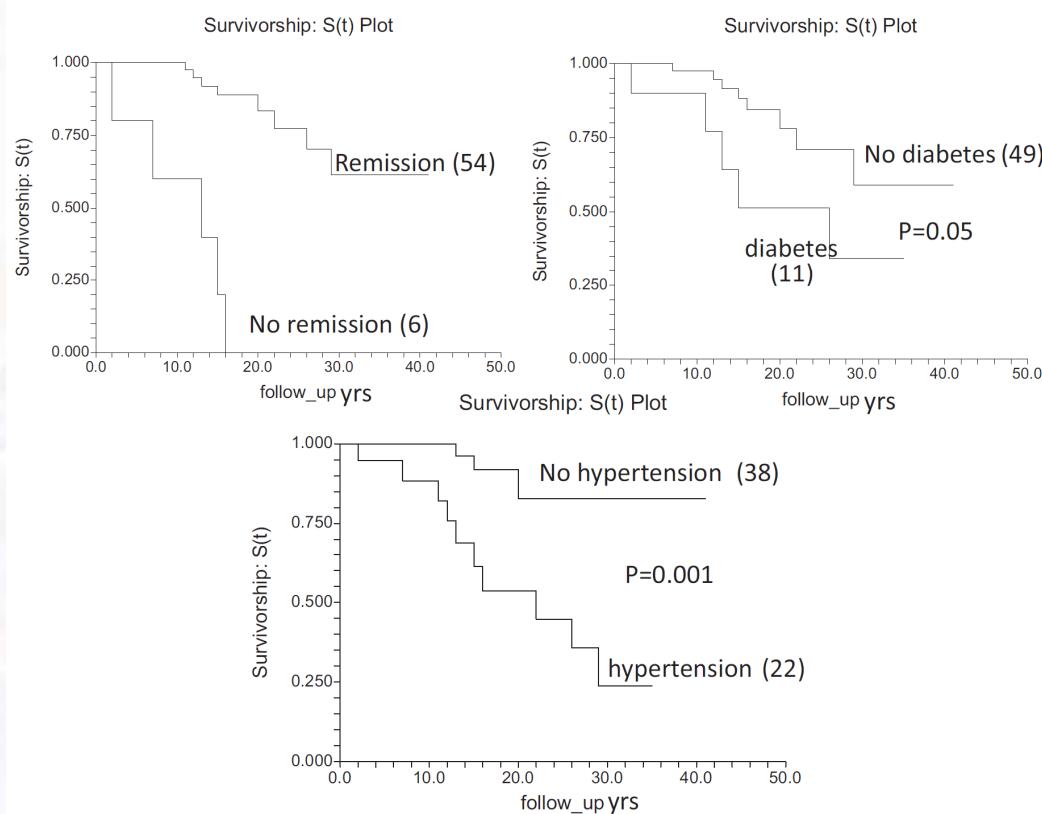
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J Clin Endocrinol Metab 86: 117-123,

TABLE 2. SMRs for Stoke Cushing's patients

	No. of deaths	Expected no. of deaths	SMR (95% CI)	P value
Overall	13	2.7	4.8 (2.8-8.3)	<0.0001
Vascular	9	0.65	13.8 (7.2-26.5)	<0.0001
Remission	8	2.4	3.3 (1.7-6.7)	0.0006
Persistent	5	0.3	16.0 (6.7-38.4)	<0.0001
Diagnosis before 1985	10	1.6	6.1 (3.3-11.4)	<0.0001
Diagnosis after 1985	3	1.1	2.8 (0.9-8.6)	0.076



Predictors of Mortality and Long-term Outcomes in Treated Cushing's Disease: A Study of 346 Patients

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- 346 pax
- 6.3 años seguimiento
- Mortalidad total 9%
(independiente de GC y
dependiente depresión)
- 30 eventos CV (incluye
ETEV).

Table 4. Predictors of Cardiovascular Events in the Total Cohort

Variable	HR (95% CI)	P Value
Age at diagnosis	1.056 (1.026–1.088)	0.000
Male sex	2.969 (1.026–6.215)	0.004
Depression	2.315 (1.057–5.068)	0.036
Diabetes	2.645 (1.270–5.505)	0.009

Multivariate regression model included age at diagnosis, sex, preoperative diabetes, and depression. Total number of subjects = 345, number of events = 30.



Risk of cardiovascular events in people prescribed glucocorticoids with iatrogenic Cushing's syndrome: cohort study

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10-12 NOVIEMBRE 2011
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Table 5 | Adjusted hazard ratios (95% confidence intervals) of cardiovascular events in patients with iatrogenic Cushing's syndrome

Cardiovascular events	Compared with patients without iatrogenic Cushing's syndrome (n=3231)				Compared with patients not prescribed glucocorticoids (n=3282)			
	Crude hazard ratio (95% CI)	P value	Adjusted hazard ratio* (95% CI)	P value	Crude hazard ratio (95% CI)	P value	Adjusted hazard ratio† (95% CI)	P value
All (n=341)	2.33 (1.73 to 3.14)	<0.001	2.74 (2.06 to 3.62)	<0.001	3.70 (2.63 to 5.22)	<0.001	4.16 (2.98 to 5.82)	<0.001
Coronary heart disease (n=177)	1.96 (1.29 to 2.97)	0.002	2.27 (1.48 to 3.47)	<0.001	2.97 (1.91 to 4.64)	<0.001	2.68 (1.62 to 4.44)	<0.001
Cerebrovascular event (n=63)	2.26 (1.06 to 4.81)	0.02	2.23 (0.96 to 5.17)	0.07	1.79 (0.85 to 3.76)	0.12	2.14 (0.97 to 4.73)	0.06
Heart failure (n=101)	2.97 (1.82 to 4.84)	<0.001	3.77 (2.41 to 5.90)	<0.001	9.74 (4.98 to 19.06)	<0.001	13.31 (7.24 to 24.51)	<0.001

*Adjusted for age, sex, initial dosage of glucocorticoids and duration of use, underlying disease, smoking status, and use of aspirin, oral anticoagulants, diabetes drugs, antihypertensive drugs, and cholesterol lowering drugs.

†Adjusted for age, sex, underlying disease, smoking status, and use of aspirin, oral anticoagulants, diabetes drugs, antihypertensive drugs, and cholesterol lowering drugs.

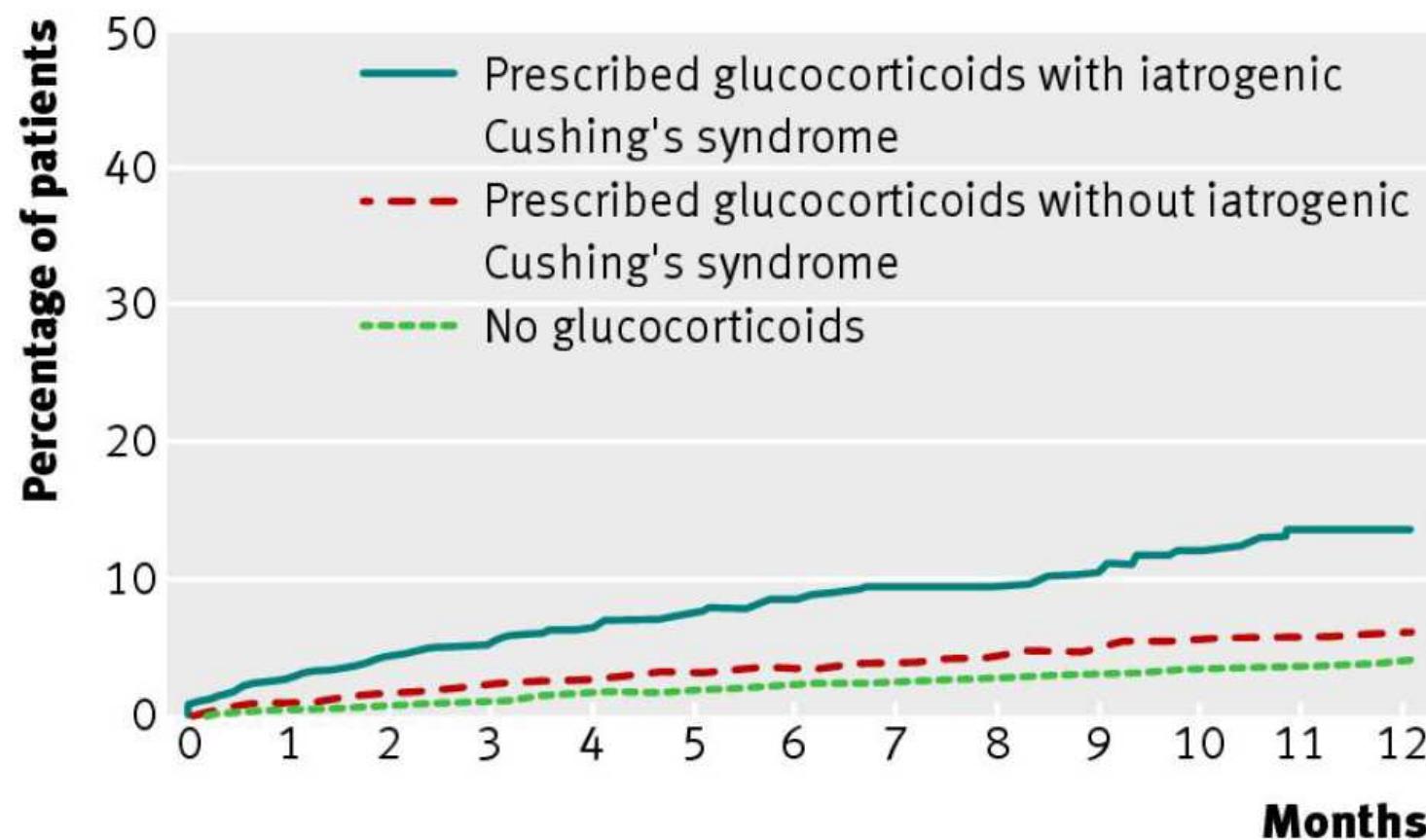


Risk of cardiovascular events in people prescribed glucocorticoids with iatrogenic Cushing's syndrome: cohort study

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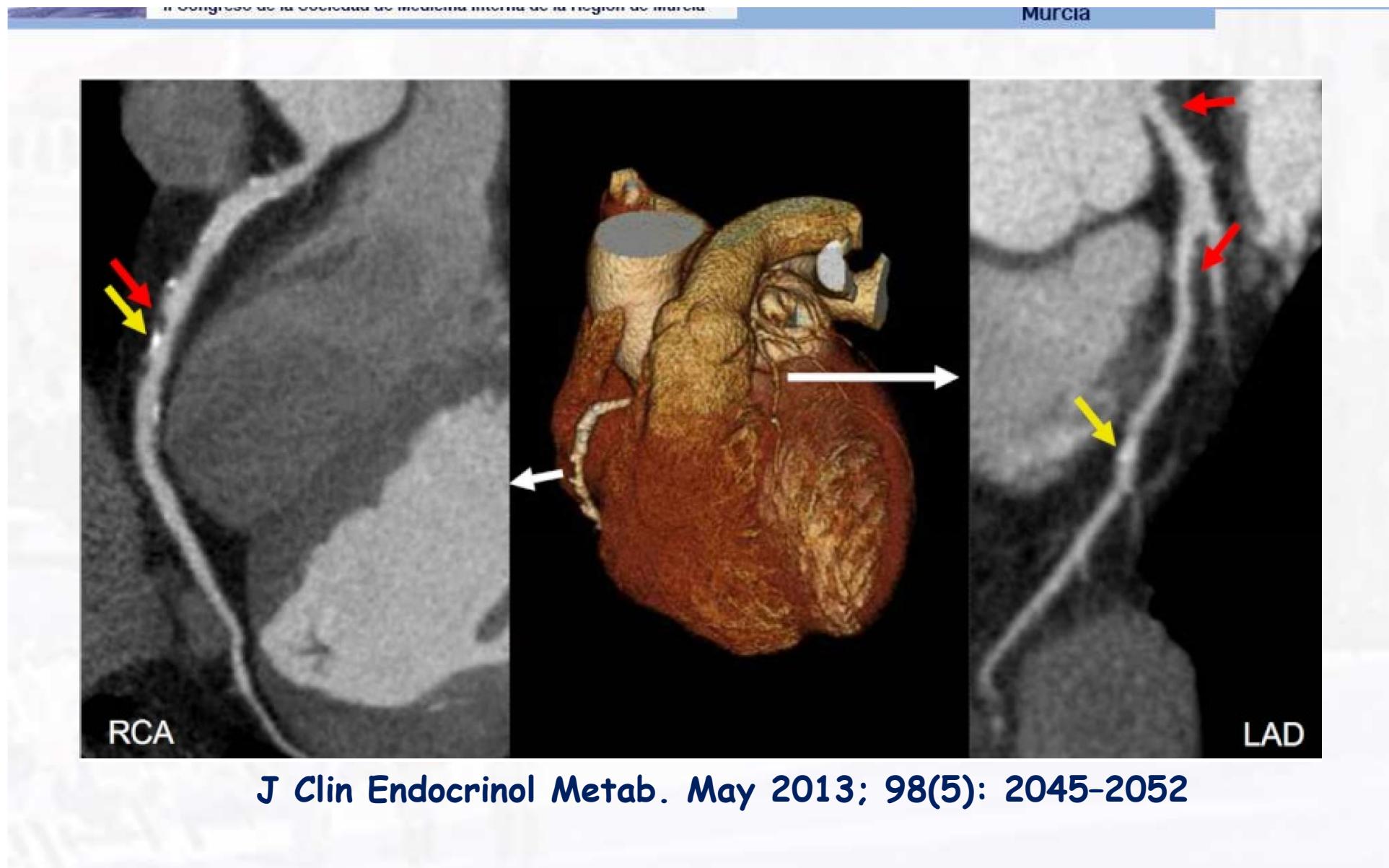
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Kaplan-Meier survival curves showing cumulative incidence of cardiovascular events over time in the three study groups



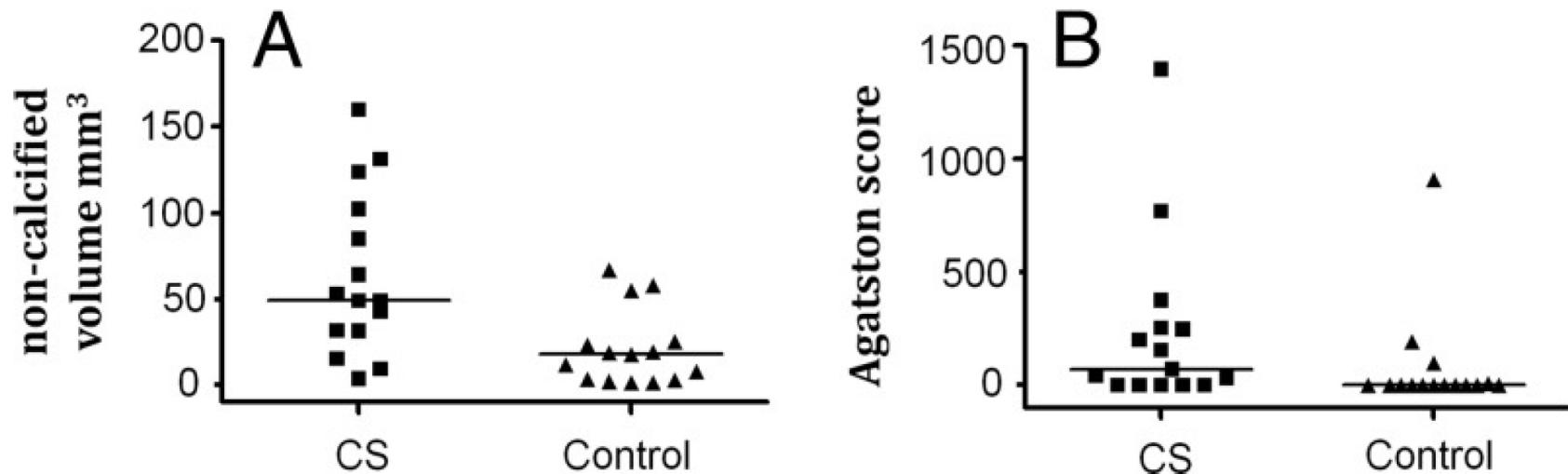
BMJ 2012;345:e4928

Hypercortisolism Is Associated With Increased Coronary Arterial Atherosclerosis



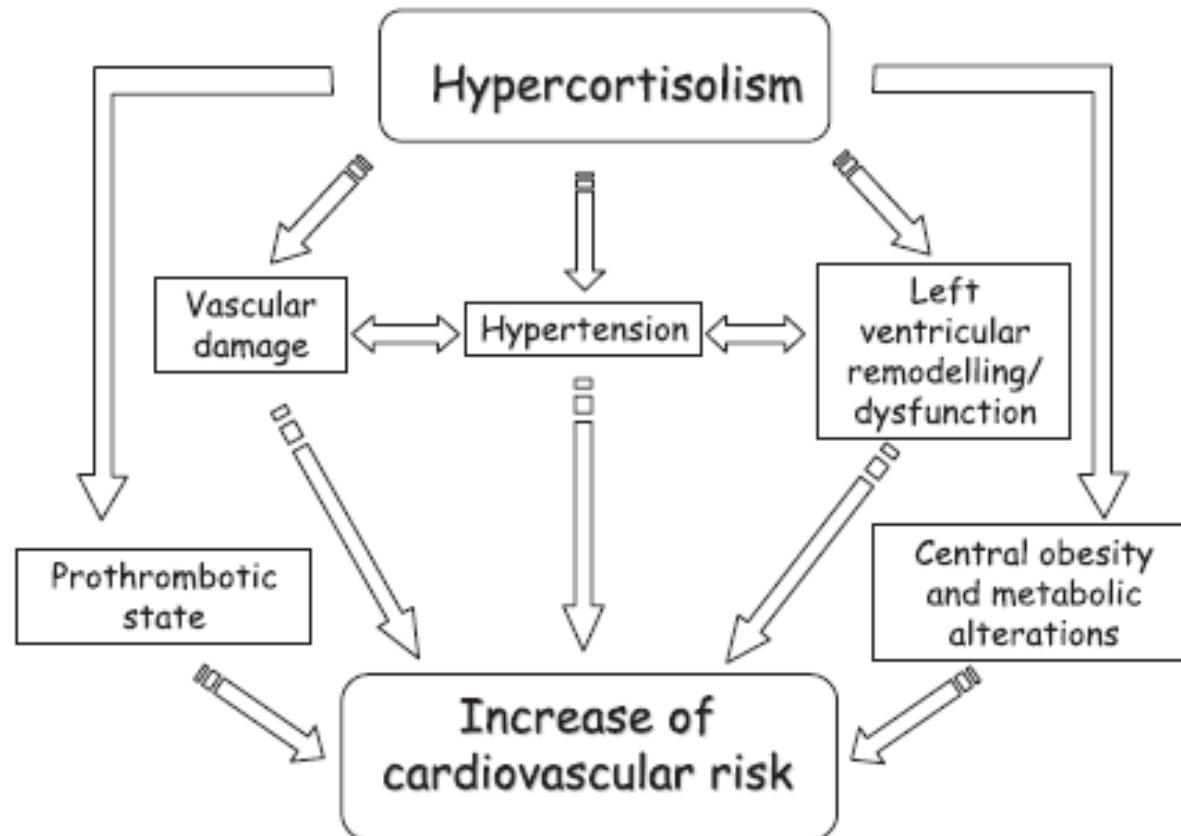
J Clin Endocrinol Metab. May 2013; 98(5): 2045-2052

Hypercortisolism Is Associated With Increased Coronary Arterial Atherosclerosis



Scatter plots in patients with CS and age-, sex-, and BMI-matched controls of (A) noncalcified plaque volume ($P < .001$ between groups) and (B) Agatston score, a measure of calcified plaque volume ($P < .05$ between groups). The horizontal lines represent the median value for each group.

Mechanisms of increased cardiovascular risk mediated by hypercortisolism

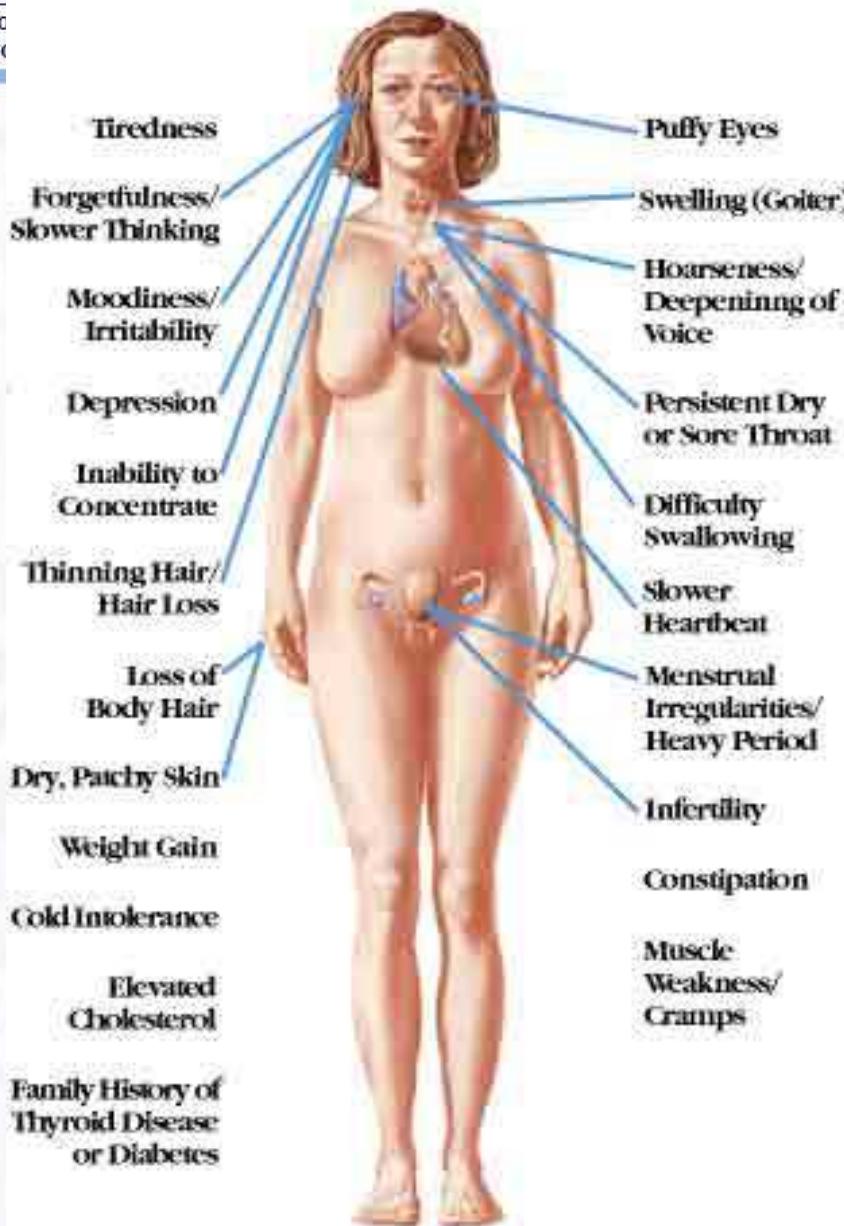


Clinical Endocrinology (2009) 71, 768-771



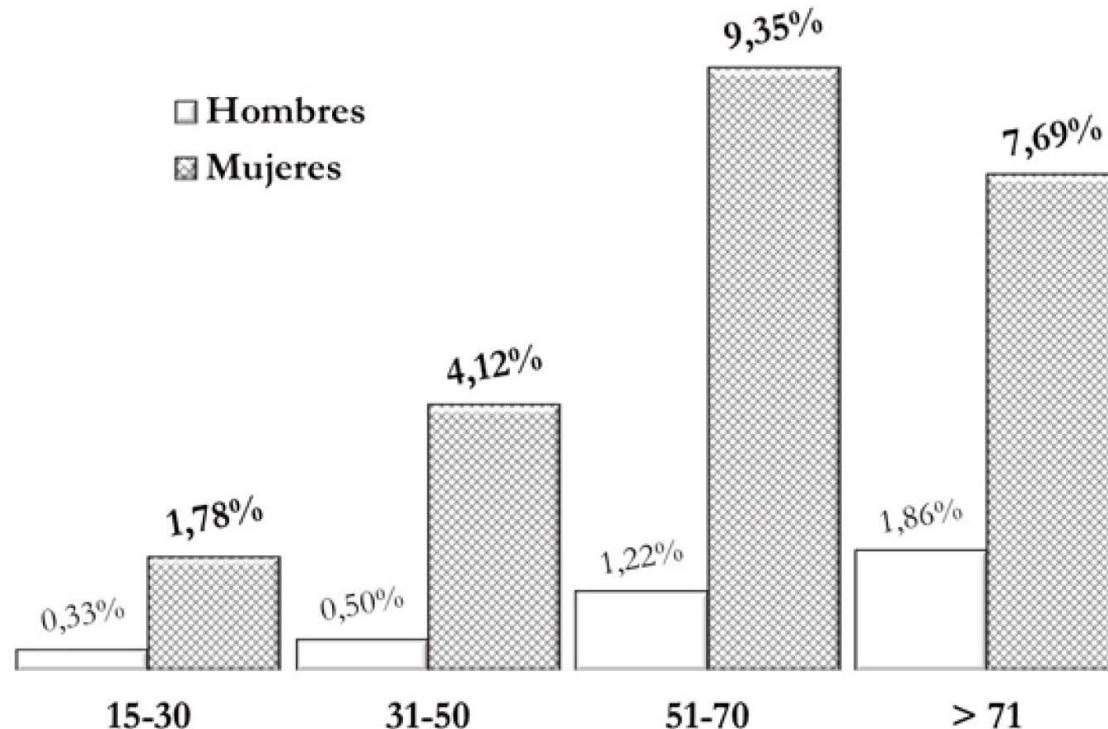
Signs and Symptoms of **HYPOTHYROIDISM**

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Hipotiroidismo

Prevalencia de HT en la provincia de Cádiz para la población mayor de 15 años, estratificada por sexo y grupos de edad, estimada únicamente por el Registro de Pacientes Tratados (RPT)





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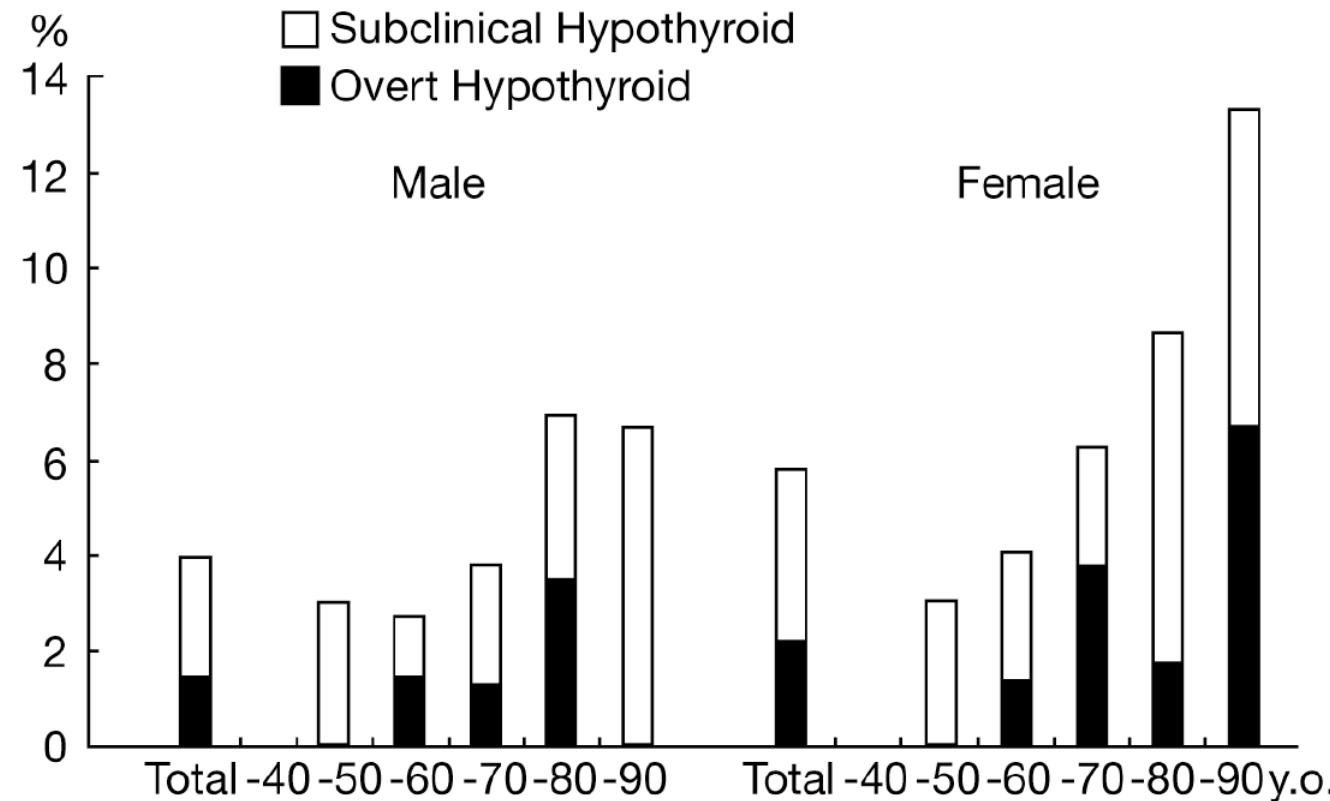


Fig. 4 The prevalence of primary hypothyroidism in each age group.

Subclinical Hypothyroidism and the Risk of Coronary Heart Disease: A Meta-Analysis

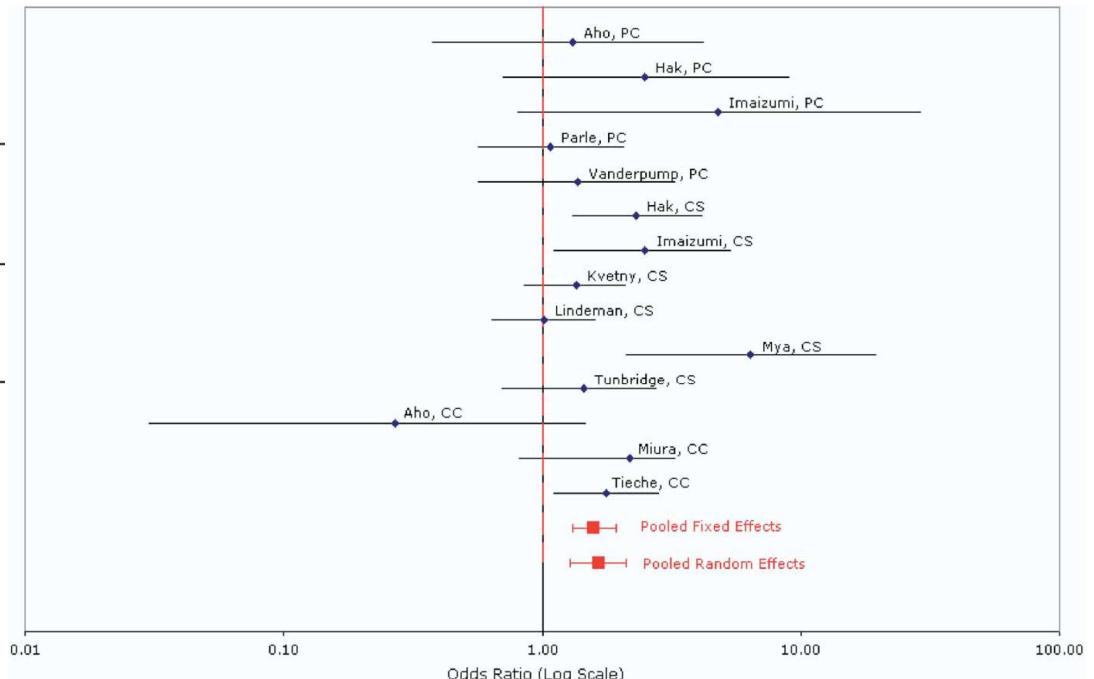
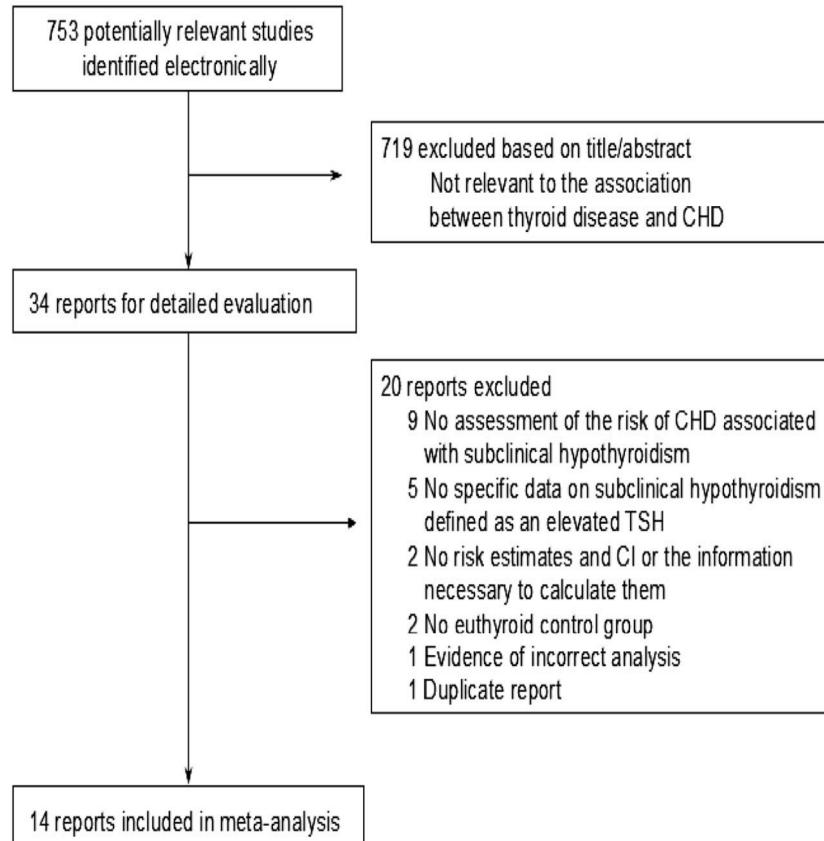


Figure 1 Forest plot of odds ratios (ORs) of coronary heart disease (CHD) associated with subclinical hypothyroidism. ORs (diamonds) and 95% confidence intervals (CIs) (horizontal lines) of the effect of subclinical hypothyroidism on the risk of CHD. CC = case-control study; CS = cross-sectional study; PC = prospective cohort study.

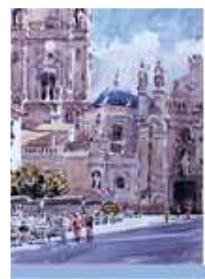
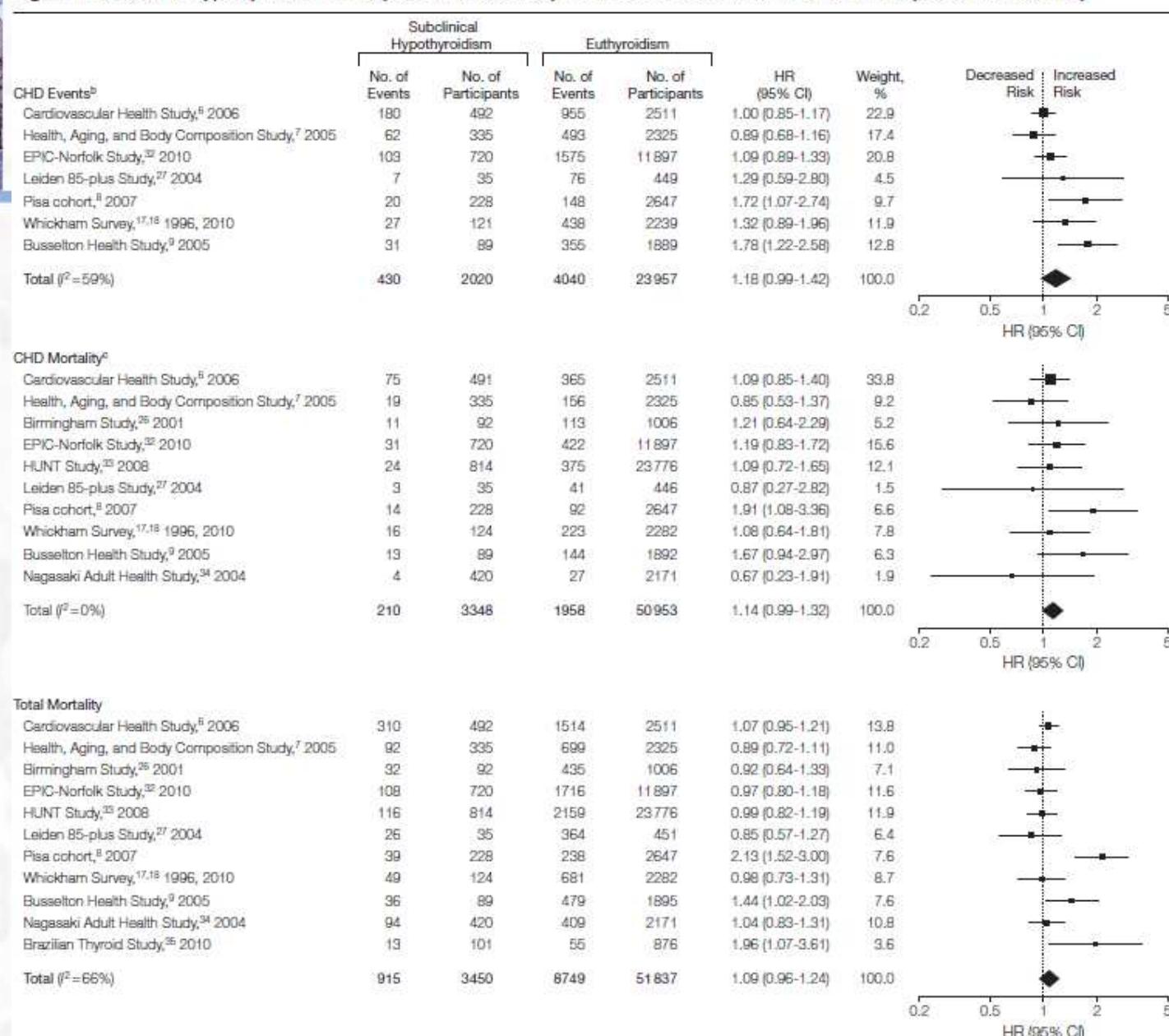


Figure 1. Subclinical Hypothyroidism vs Euthyroidism for Coronary Heart Disease (CHD) Events, CHD Mortality, and Total Mortality^a





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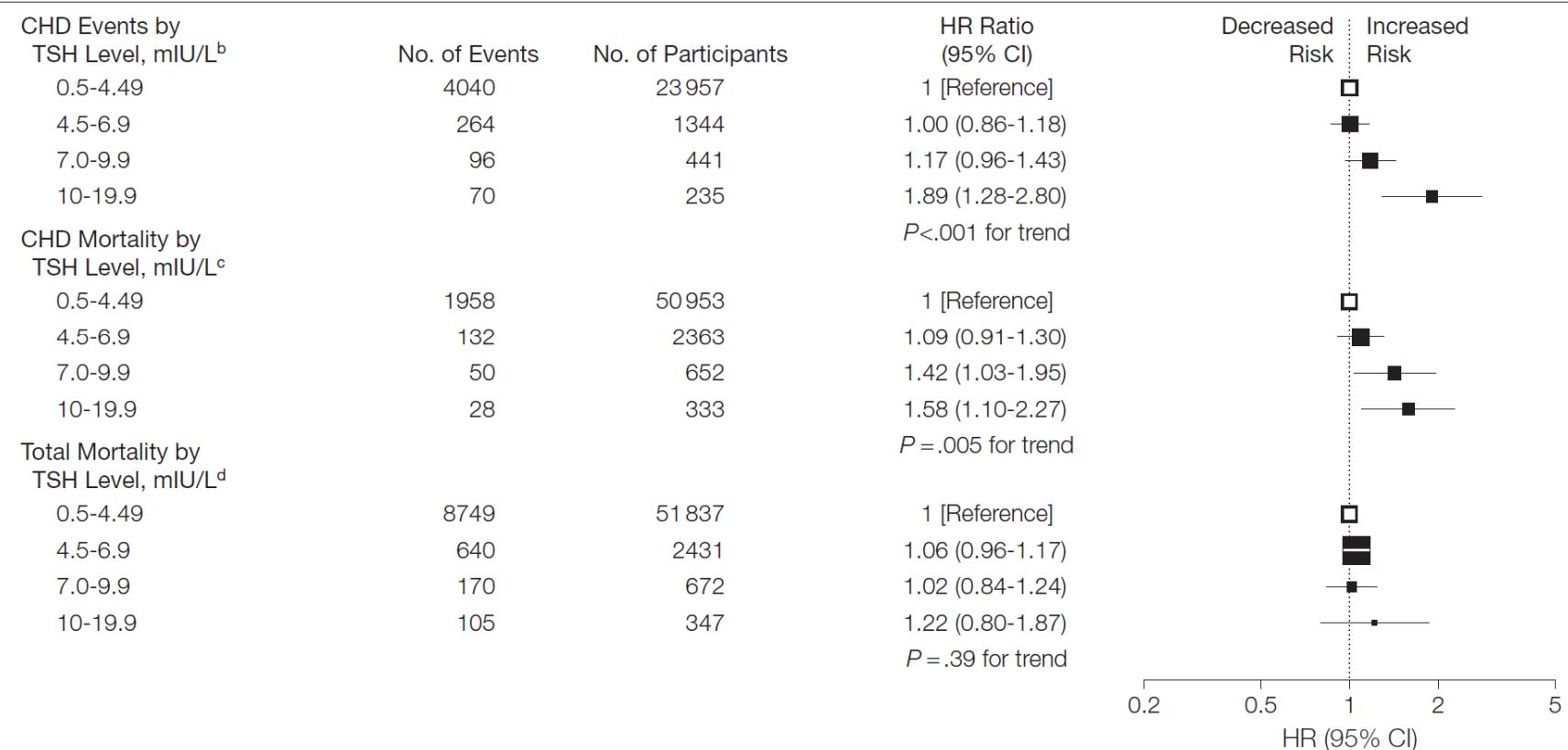
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Figure 2. Hazard Ratios (HRs) for Coronary Heart Disease (CHD) Events, CHD Mortality, and Total Mortality According to Elevated Thyroid-Stimulating Hormone (TSH) Categories and Subclinical Hypothyroidism Stratified by Age vs Euthyroidism^a





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Murcia

Thyroid hormone replacement for subclinical hypothyroidism (Review)

Villar HCCE, Saconato H, Valente O, Atallah ÁN

Authors' conclusions

In current RCTs, levothyroxine replacement therapy for subclinical hypothyroidism did not result in improved survival or decreased cardiovascular morbidity. Data on health-related quality of life and symptoms did not demonstrate significant differences between intervention groups. Some evidence indicates that levothyroxine replacement improves some parameters of lipid profiles and left ventricular function.





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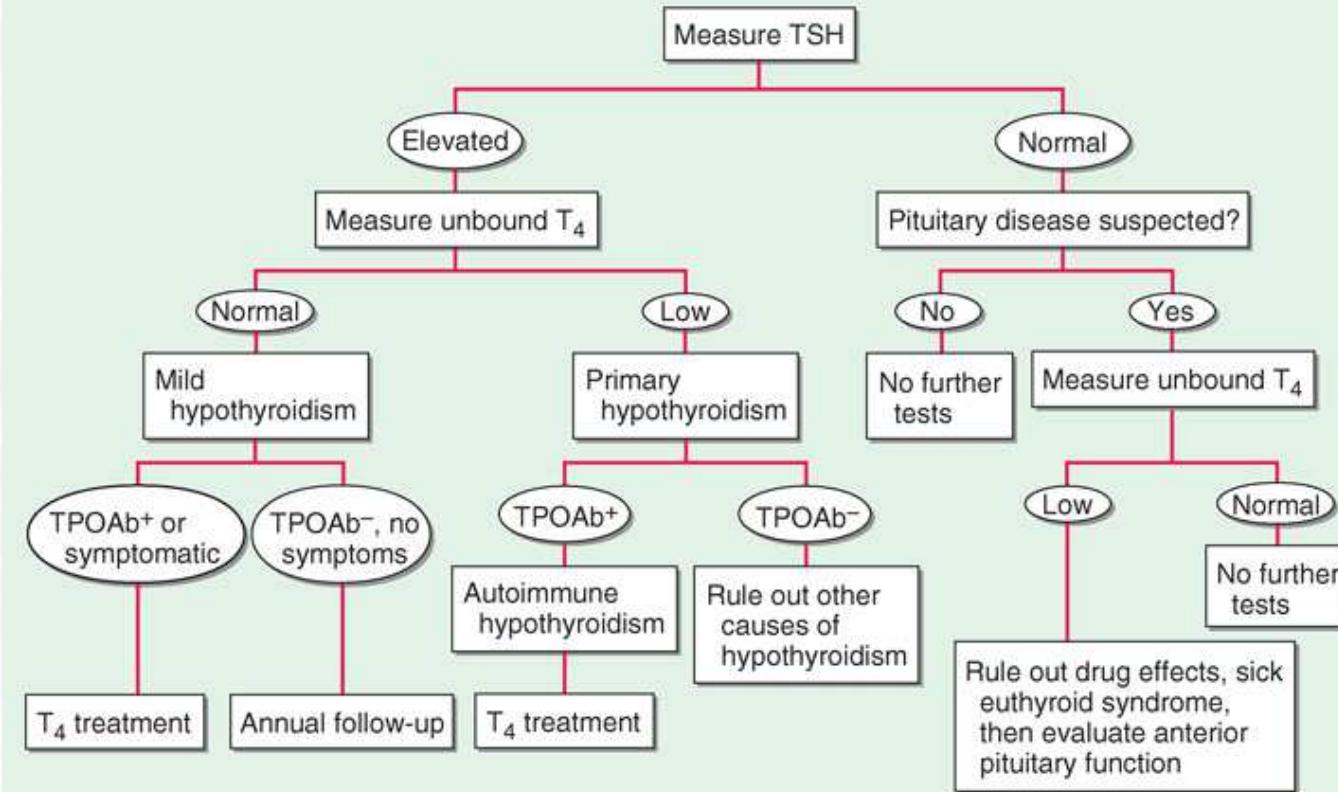
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EVALUATION OF HYPOTHYROIDISM



Source: Longo DL, Fauci AS, Kasper DL, Hauser SL, Jameson JL, Loscalzo J: *Harrison's Principles of Internal Medicine*, 18th Edition: www.accessmedicine.com

Conclusiones

- Las enfermedades aquí incluidas se acompañan de un incremento de la morbilidad y mortalidad coronarias
- La expresión clínica de las enfermedades endocrinas va cambiando a expensas de las formas subclínicas
- El tratamiento “curativo” de la enfermedad no siempre revierte los cambios funcionales o estructurales.

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