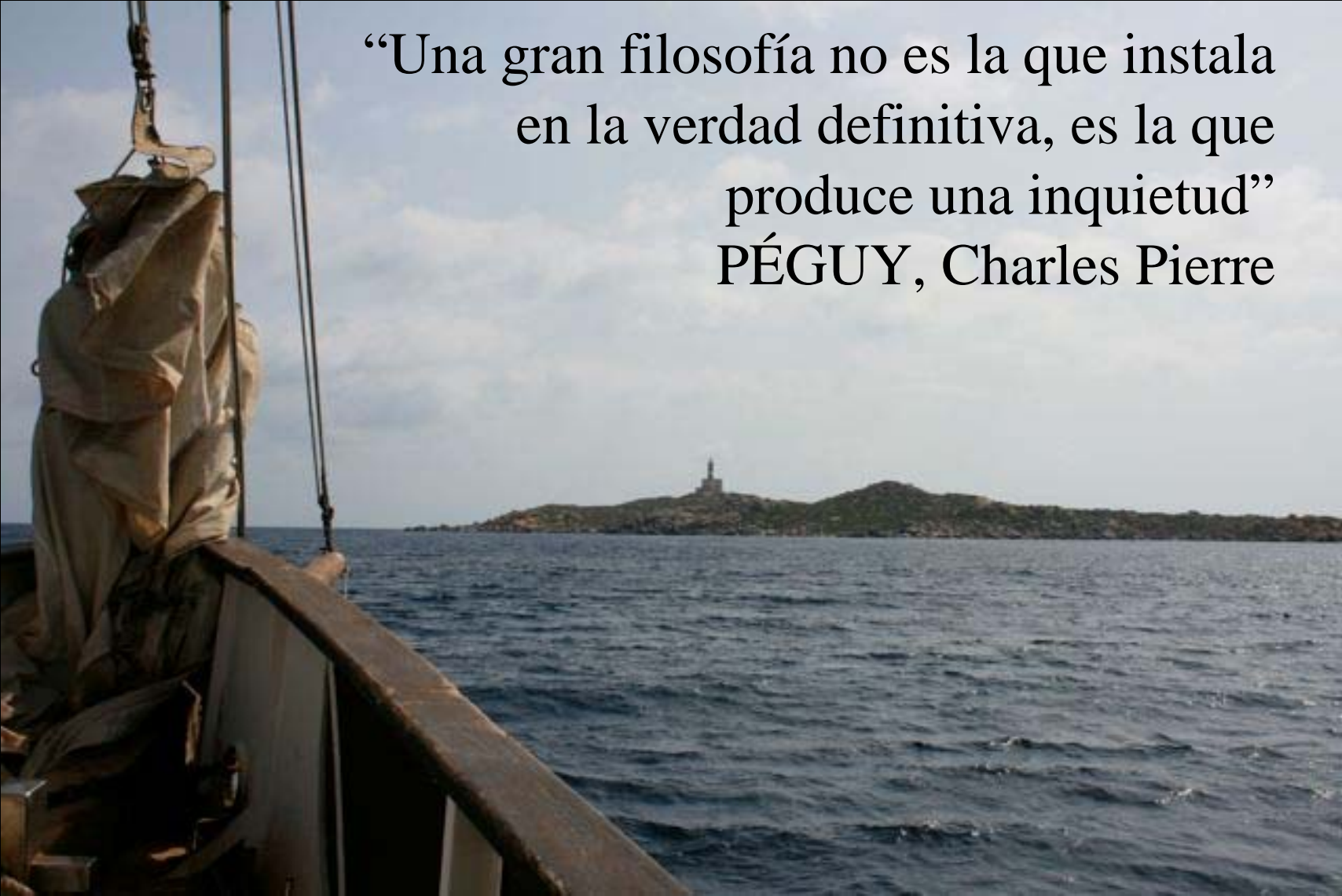


III Reunión Nacional de Actualización en Enfermedades
Autoinmunes Sistémicas para Residentes

Profilaxis de la tuberculosis en el paciente inmunodeprimido: ¿cuándo?

Juan José Ríos Blanco



“Una gran filosofía no es la que instala
en la verdad definitiva, es la que
produce una inquietud”
PÉGUY, Charles Pierre



Clinical Practice Guidelines



American Thoracic Society

Targeted Tuberculin Testing and Treatment of Latent Tuberculosis Infections

THIS OFFICIAL STATEMENT OF THE AMERICAN THORACIC SOCIETY WAS ADOPTED BY THE ATS BOARD OF DIRECTORS, JULY 1999. THIS IS A JOINT STATEMENT OF THE AMERICAN THORACIC SOCIETY (ATS) AND THE CENTERS FOR DISEASE CONTROL AND PREVENTION (CDC). THIS STATEMENT WAS ENDORSED BY THE COUNCIL OF THE INFECTIOUS DISEASES SOCIETY OF AMERICA (ILSA), SEPTEMBER 1999, AND THE SECTIONS OF THIS STATEMENT AS IT RELATES TO INFANTS AND CHILDREN WERE ENDORSED BY THE AMERICAN ACADEMY OF PEDIATRICS (AAP), AUGUST 1999.

Although the terms “preventive therapy” and “chemoprophylaxis” have been used for decades, they have also been confusing. “Preventive therapy” has referred to the use of a simple regimen (usually isoniazid) to prevent the development of active TB disease in persons known or likely to be infected with *M. tuberculosis*, but it rarely results in true primary prevention (i.e., prevention of infection in persons exposed to persons with infectious TB). To describe the intended intervention more accurately, this report uses the terminology “treatment of LTBI” rather than “preventive therapy” or “chemoprophylaxis.” This change in nomenclature will hopefully promote greater understanding of the concept for both patients and providers, resulting in more widespread implementation of this essential TB control strategy.

TABLE 7

CRITERIA FOR TUBERCULIN POSITIVITY, BY RISK GROUP

Reaction \geq 5 mm of Induration	Reaction \geq 10 mm of Induration
Human immunodeficiency virus (HIV)-positive persons	Recent immigrants (i.e., within the last 5 yr) from high-prevalence countries
Recent contacts of tuberculosis (TB) case patients	Injection drug users
Fibrotic changes on chest radiograph consistent with prior TB	Residents and employees† of the following high-risk congregate settings: prisons and jails, nursing homes and other long-term facilities for the elderly, hospitals and other health care facilities, residential facilities for patients with acquired immunodeficiency syndrome (AIDS), and homeless shelters
Patients with organ transplants and other immunosuppressed patients (receiving the equivalent of \geq 15 mg/d of prednisone for 1 mo or more)*	Mycobacteriology laboratory personnel
	Persons with the following clinical conditions that place them at high risk: HIV infection, diabetes mellitus, chronic renal failure, some hematologic disorders (e.g., leukemias and lymphomas), other specific malignancies (e.g., carcinoma of the head or neck and lung), weight loss of \geq 10% of ideal body weight, gastrectomy, and jejunioileal bypass
	Children younger than 4 yr of age or infants, children, and adolescents exposed to adults at high-risk

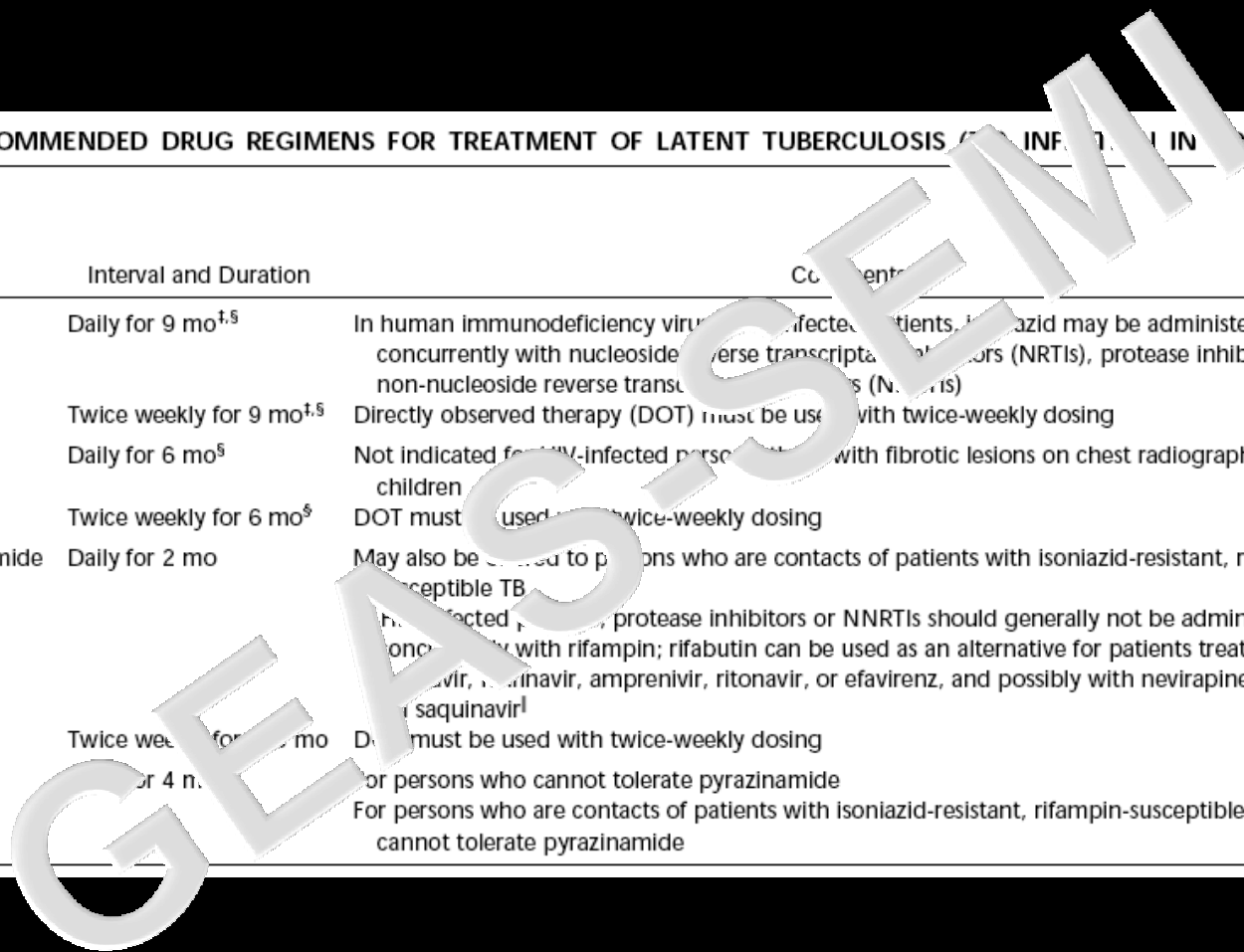
* Risk of TB in patients treated with corticosteroids increases with higher dose and longer duration.

† For persons who are otherwise at low risk and are tested at the start of employment, a reaction of \geq 15 mm induration is considered positive.

Source: Adapted from Centers for Disease Control and Prevention. Screening for tuberculosis and tuberculosis infection in high-risk populations. *M.M.W.R.* 1995;44(No. RR-11):19-34.

RECOMMENDED DRUG REGIMENS FOR TREATMENT OF LATENT TUBERCULOSIS (LTB) IN HIV-1 POSITIVE ADULTS

Drug	Interval and Duration	Comments	Rating* (Evidence) [†]	
			HIV ⁻	HIV ⁺
Isoniazid	Daily for 9 mo ^{4,5}	In human immunodeficiency virus (HIV)-infected patients, isoniazid may be administered concurrently with nucleoside reverse transcriptase inhibitors (NRTIs), protease inhibitors, or non-nucleoside reverse transcriptase inhibitors (NNRTIs)	A (II)	A (II)
Isoniazid	Twice weekly for 9 mo ^{4,5}	Directly observed therapy (DOT) must be used with twice-weekly dosing	B (II)	B (II)
Isoniazid	Daily for 6 mo ⁵	Not indicated for HIV-infected persons with fibrotic lesions on chest radiographs, or children	B (I)	C (I)
Isoniazid	Twice weekly for 6 mo ⁵	DOT must be used with twice-weekly dosing	B (II)	C (I)
Rifampin plus pyrazinamide	Daily for 2 mo	May also be considered to persons who are contacts of patients with isoniazid-resistant, rifampin-susceptible TB	B (II)	A (I)
Rifampin plus pyrazinamide	Twice weekly for 2 mo	DOT must be used with twice-weekly dosing	C (II)	C (I)
Rifampin	Daily for 4 mo	For persons who cannot tolerate pyrazinamide	B (II)	B (III)
Rifampin	Twice weekly for 4 mo	For persons who are contacts of patients with isoniazid-resistant, rifampin-susceptible TB who cannot tolerate pyrazinamide		





*Normativa SEPAR:
Diagnóstico
y tratamiento
de la tuberculosis*

RECOMENDACIONES SEPAR



Tabla 1. Indicaciones de la prueba de la tuberculina

Convivientes y contactos íntimos de pacientes tuberculosos
Personas cuya radiografía de tórax presente imágenes indicativas de tuberculosis inactiva
Personas con sospecha clínica y/o radiológica de presentar enfermedad tuberculosa
Personas que si están infectadas tienen un especial riesgo de desarrollar enfermedad tuberculosa
Infección por el virus de la inmunodeficiencia humana
Adicción a drogas por vía parenteral
Marginación social
Enfermedades inmunodepresoras: leishmaniasis, sífilis, toxoplasmosis, neosporosis, citomegalovirus, hepatitis, sarcoidosis, linfoma y otras neoplasias
Tratamiento inmunodepresor prolongado, tratamiento contra el factor de necrosis tumoral alfa y otros fármacos a largo plazo
Personas que si están infectadas constituyen un riesgo social y epidemiológico en caso de desarrollar tuberculosis activa
Cuidadores de guarderías infantiles
Personas con discapacidad
Personal sanitario
Personal de prisiones
Estudios epidemiológicos y control de programas antituberculosos

GEASIS-SEMI

20-10-03

$\geq 5\text{mm}$



Tabla 9. Tratamiento de la infección tuberculosa

Indicaciones (A)

- Infección reciente (contactos, conversión tuberculínica)
- Coinfectados por el VIH
- Lesiones radiológicas de tuberculosis inactiva no tratada

Quimioprofilaxis primaria (tratamiento de expuestos sin infección)

- Niños menores de 5 años (D)
- Infección por el VIH (D)
- Niños y adolescentes (valoración individual) (D)

Inmunodeprimidos

Tipo de tratamiento

1. Hidracidas durante 6 meses con pauta estándar (A)
 2. Hidracidas durante 4 meses en pacientes con infección por el VIH (B)
 3. Rifampicina e hidracidas durante 3 meses (B)
 4. Rifampicina e hidracidas durante 3 meses con lesiones residuales en la radiografía de tórax (B)
 5. Rifampicina e hidracidas durante 3 meses con pauta alternativa a hidracidas durante 6 meses
 6. Rifampicina durante 4 meses, en pacientes sin infección por el VIH (B), en pacientes con infección por el VIH (D)
- En resistencia a hidracidas

VIH: virus de la inmunodeficiencia humana

Indicaciones de quimioprofilaxis (tratamiento de la infección tuberculosa)

QPP (negativos para la tuberculina)

Jóvenes contactos íntimos de bacilíferos

Inmunodeprimidos contactos de bacilíferos

Personas de cualquier edad, contactos íntimos de bacilíferos en las microepidemias

QPS o TIT (positivos para la tuberculina)

Indicaciones prioritarias

Infectados por el VIH

Convertidores tuberculínicos*

Miembros de microepidemias de cualquier edad

Silicosis

Imágenes fibróticas residuales no tratadas**

Menores de 35 años, contactos de enfermos bacilíferos

Cualquier infección menor de 6 meses

Pacientes en lista de espera de trasplantes

Utilización de inmunoglobulinas o anticuerpos monoclonales anti- α

Indicaciones para tratar actualmente

Mayor de 2 años, contactos de enfermos bacilíferos

Toxicomanías, incluido alcoholismo

Trastorno oncológico

Neoplasias

Tratamientos prolongados con corticoides

o inmunodepresores

Insuficiencia renal crónica. Hemodiálisis

Desnutrición: gastrectomía, síndromes de malabsorción y derivación intestinal

Riesgo profesional: docentes y guarderías, sanitarios, trabajadores con grupos de riesgo (prisiones, asilos, centros de toxicómanos, etc.)

Riesgo social: asilados, reclusos, albergues, psiquiátricos, etc.

Inmigrantes de bajo nivel económico

Estudio clínico de la tuberculosis en inmunodeprimidos

Valencia ME, Gil A, Díaz A, Torres A, Lavilla P, Pintado V, López D, ... Vázquez JJ
Rev Clin Esp 1989; 184: 352-356

- neoplasias
- ADVP
- TB
- No TB



Incidencia:
429/100 000
x10

4 TB extrapulmonar / 2 fallecieron

Incidencia y características de la tuberculosis en pacientes con enfermedades reumáticas autoinmunes

C. Vadillo Font, C. Hernández-García, E. Pato, I. C. Morado, M. Salido, E. Júdez, P. Macián, B. Fernández-Gutiérrez, L. Abásolo y J. A. Jover

Servicio de Reumatología. Hospital Clínico San Carlos. Madrid

Densidad de incidencia de tuberculosis por diagnóstico

Diagnóstico	Pacientes	Casos TB	Pacientes-año	Densidad de incidencia*	IC 95%
ACG	88	1	3.172	367	(9-2.031)
PMR	79	1	178	565	(14-3.107)
Otras vasculitis	87	0	930	930	(113-3.320)
Artritis reumatoide	1.016	8	3.460	231	(100-455)
Espondiloartropatías	892	0	2.275	—	(<162)***
Lupus eritematoso sistémico	158	5	466	645	(133-1.870)
Esclerodermia	42	0	124	—	(<2.931)***
Síndrome de Sjögren	76	0	157	—	(<2.322)***
Miopatías inflamatorias	15	0	62	—	(<5.776)***
EMTC	21	0	106	—	(<3.420)***
Poliartritis no filiadas	29	0	738	—	(<499)***
Uveítis**	264	0	738	—	(<499)
Total	3.634	15	9.795	153	(86-252)

*Por 100.000 pacientes-año. **No se incluyen pacientes con panuveítis, uveítis intermedia y posterior. ***Intervalo de confianza del 95,5%. ACG: arteritis de células gigantes; PMR: polimialgia reumática; EMTC: enfermedad mixta del tejido conectivo.

DI población general: 31/100000 hab-año x5

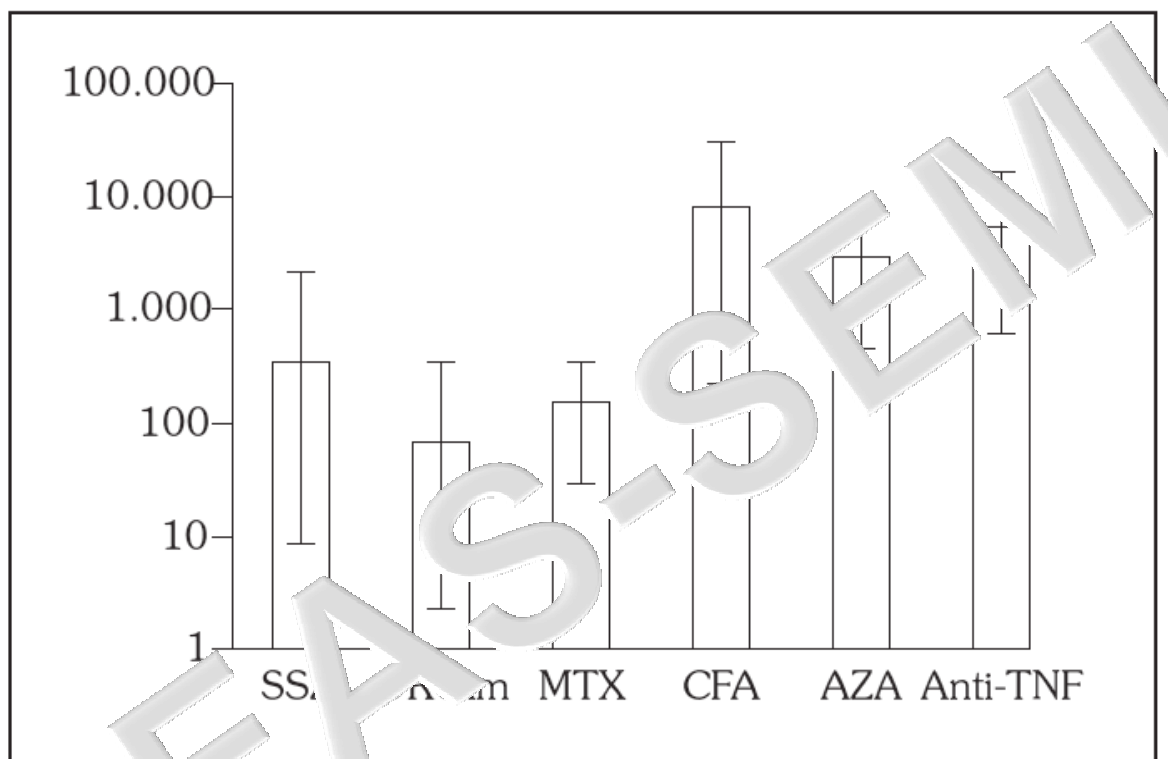


Fig. 1. Incidencia de tuberculosis (en escala logarítmica) en pacientes con artritis reumatoide tratados con diferentes fármacos modificadores de la enfermedad. SSZ: sulfasalazina; Oroim: aurotiomalato sódico; MTX: metotrexato; CFA: ciclofosfamida; AZA: azatioprina.

High risk of tuberculosis in systemic lupus erythematosus?

J-G Erdozain, G Ruiz-Irastorza*, M-V Egurbide, A Martinez-Piñero, and T Aguirre
 Department of Internal Medicine, Hospital de Cruces, University of the Basque Country, Bizkaia, Spain

1994-2003: 1063 pacientes-año (0 cases of tuberculosis) X 7

Table 1 Summary of published series of tuberculosis in patients with systemic lupus erythematosus

Series (ref.)	Study years	Cases per 100 000 patient-years (95% CI)	Extrapulmonary TB %	Mortality %
Philippines (9)	1985-1995	1461 (1102-1897)	56	14
Hong Kong (10)	1984-2001	122 (525-877)	66	7
Korea (11)	1979-2000	75 (21-131)	60	13
Korea II (12)	1990-1995	587 (215-945)	NR	5
India (13)	1989-1994	173 (173-3347)	22	NR
Singapore (14)	1975-1979	21 (160-492)	50	31
Turkey (15)	1976-1980	50 (90-232)	45	5
India II (16)	NR	2328 (1370-3694)	47	6
Malaysia (17)	1978-1983	1067 (354-2304)	0	NR
Madrid (Spain) (18)	1994-2003	645 (129-1931)	33	0
Cruces (Spain)	1994-2003	187 (39-547)	0	0

CI: confidence interval; TB: tuberculosis; NR: not reported

Tuberculosis in patients with systemic rheumatic or pulmonary diseases treated with glucocorticosteroids and the preventive role of isoniazid: a review of the available evidence

Matthew E. Falagas^{a,b,*}, Paraskevi T. Voidonikola^a, Anna G. Angelaki^a

International Journal of Antimicrobial Agents 30 (2007) 477–481

20 estudios: 16 retrospectivos, 2 prospectivos, 2 casos y controles

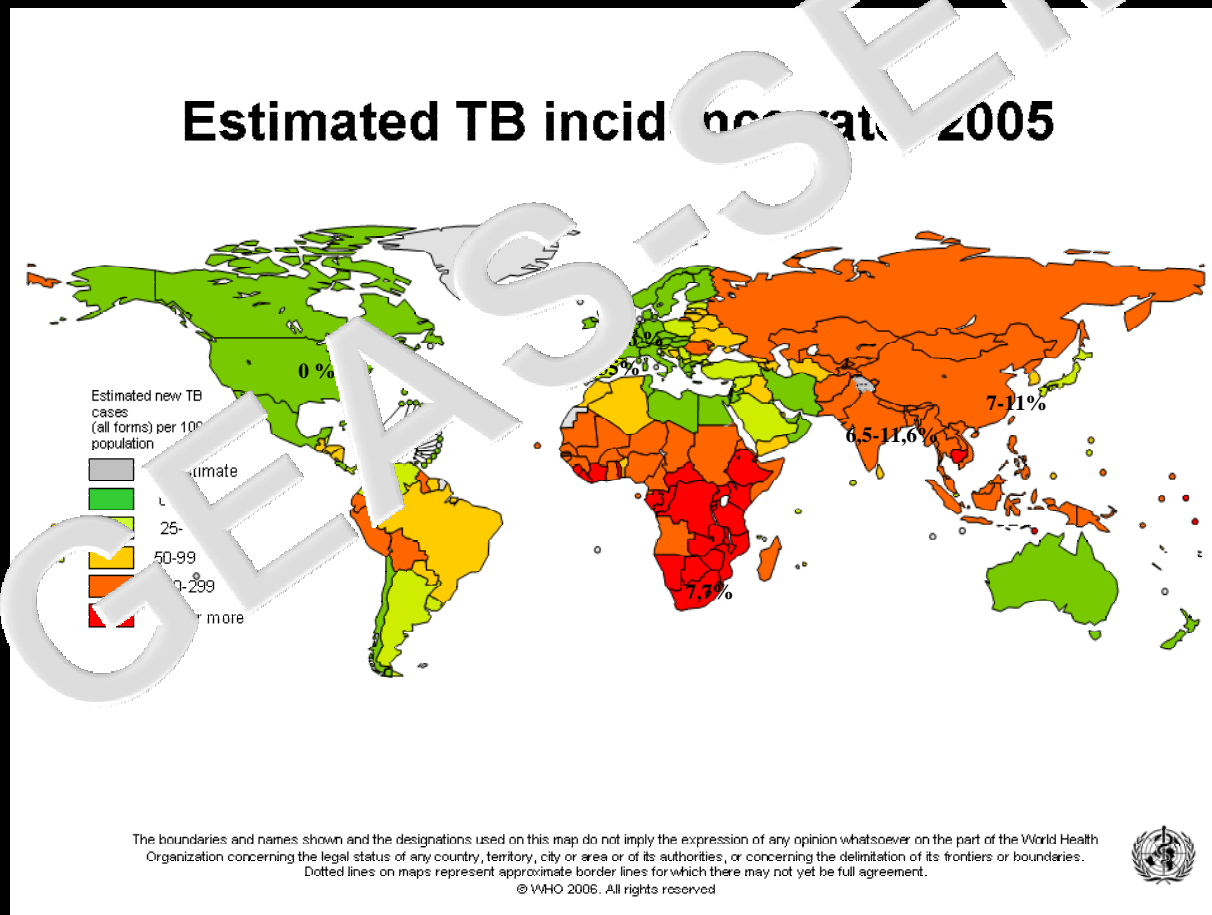


Table 3

Effect of isoniazid prophylaxis (INHP) on the development of tuberculosis (TB) in patients receiving steroids with or without other immunosuppressants for systemic rheumatic or chronic pulmonary diseases

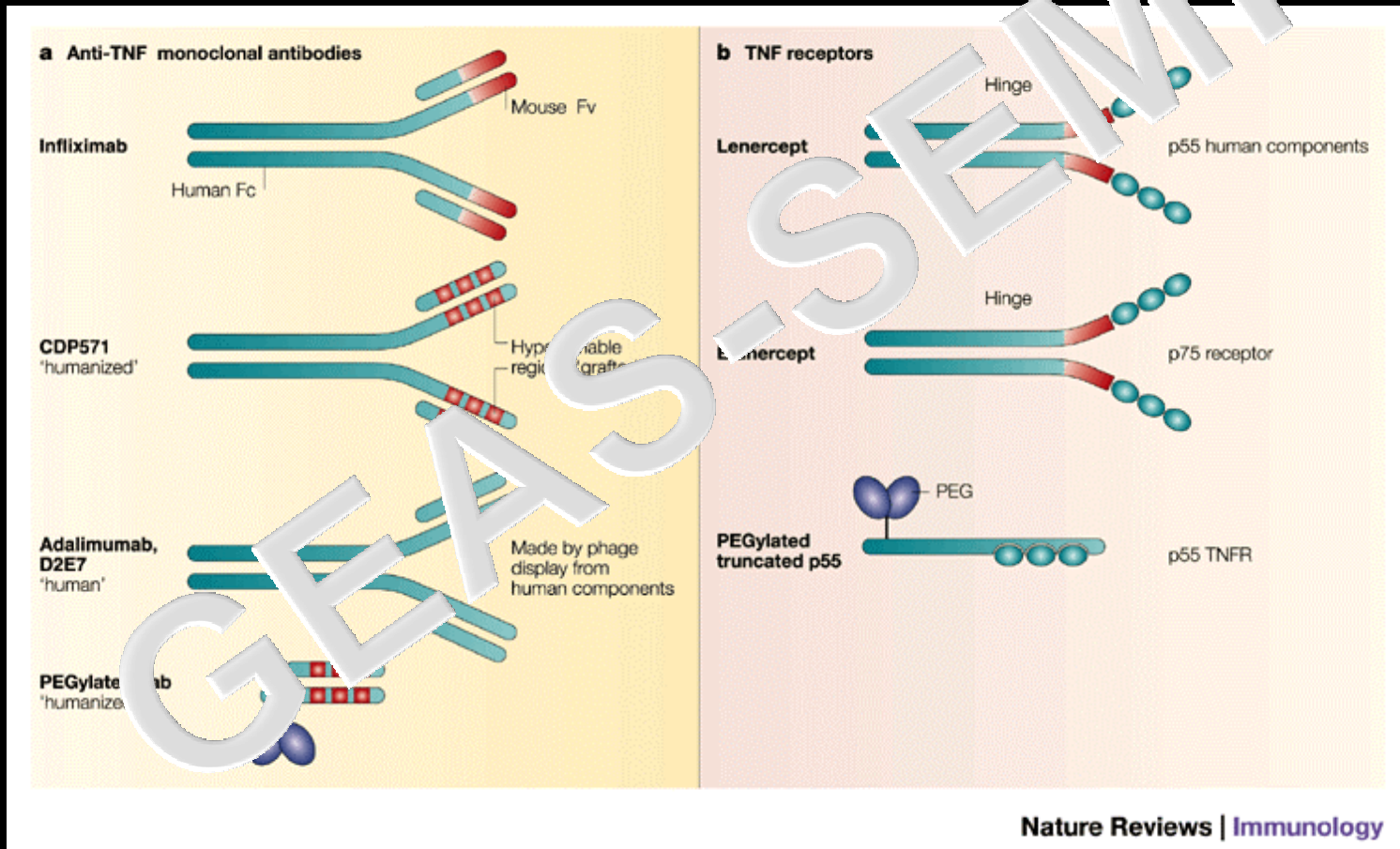
Reference (setting)	INHP	Patients with INHP	Patients without any prophylaxis	Mean \pm S.D. follow-up (years)	No. of patients with TB	Frequency of development of TB (with vs. without INHP)	Comments
Mok et al. (Hong Kong) [3]	42 patients with history of TB for which they received complete therapy (nested case-control analysis): INHP in 21 (300 mg/day INH + pyridoxine 10 mg/day for 12.4 months)	21	21 (matched) ^a	18.3 \pm 7.1 (INHP group), 18.3 \pm 7.1 (no INHP group)	2 in the INHP group vs. 4 in the no INHP group	9.5% vs. 19%	No statistically significant difference ($P = 0.66$), but small number of patients. Incidence: 0.55 vs. 1.04 per 100 patient-years (matched) ^a
Gaitonde et al. (India) [5]	INHP in 70 patients (5 mg/kg/day INH (max. 300 mg/day) + pyridoxine 10 mg/day for 1 year)	70	0		1 (in the INHP group); there was not a no INHP group in this study	1.4% vs. 11% (as estimated in a previous study of patients without INHP) ^b	82% reduction in incidence of TB with INHP
Kobashi and Matsushima (Japan) [6]	INHP in 62 patients (0.3 g/day INH for 2–6 months)	62		5	2 in the INHP group vs. 12 in the no INHP group	3.2% vs. 2.2%	No difference (but insufficient duration of INHP)
Hernandez-Cruz et al. (Mexico) [10]	INHP in 41 patients (300 mg/day INH for 0.5 year)	41	37	NR	1 in the INHP group vs. 13 in the no INHP group	2.4% vs. 35%	97% reduction of incidence of TB with INHP

NR, Not reported; ^aMatched case-control study; ^bRetrospective study.

Tratamiento esteroideo y tuberculosis

- ¿Papel otros inmunosupresores?
- Fiabilidad Mantoux.
- Distintas enfermedades, ¿distinto riesgo?
- Reactivación versus reinfección.
- Isoniacina: coste / eficacia, ¿toxicidad?

Agentes anti TNF y tuberculosis



Registro BIOBADASER

ARTHRITIS RHEUMATOIDE
Vol. 48, No. 1, p. 2122-2127

- 17/1578 infliximab (0 etanercept).

Año	Incidencia 100000 hab-año	RR población general	RR población AR pre anti-TNF (EMECAR)
2000	1812	90,1 %	19,9 %
2001	1113	53 %	11,7 %

65 % extrapulmonares

Table 2. Recommendations of the Spanish Health Authorities and the Spanish Society of Rheumatology regarding the management of TB risk in RA patients who are to undergo treatment with tumor necrosis factor inhibitors*

Patients in any of the following categories should be treated for **9 months** with 5 mg/kg body weight (up to a maximum of 300 mg) of isoniazid daily.

History of untreated or previously treated TB, or exposure to an active case of TB

Chest radiograph showing residual changes indicative of prior TB infection

Reaction **≥5 mm** in diameter on PPD skin testing or on 2-step testing procedure (when initial PPD result is <5 mm in diameter), with an interval of 7–10 days between steps

Registro BIOBADASER

ARTHRITIS & RHEUMATISM

Vol. 52, No. 6, June 2010, pp 1766–1772

Isoniacida: 324/385 (84 %)

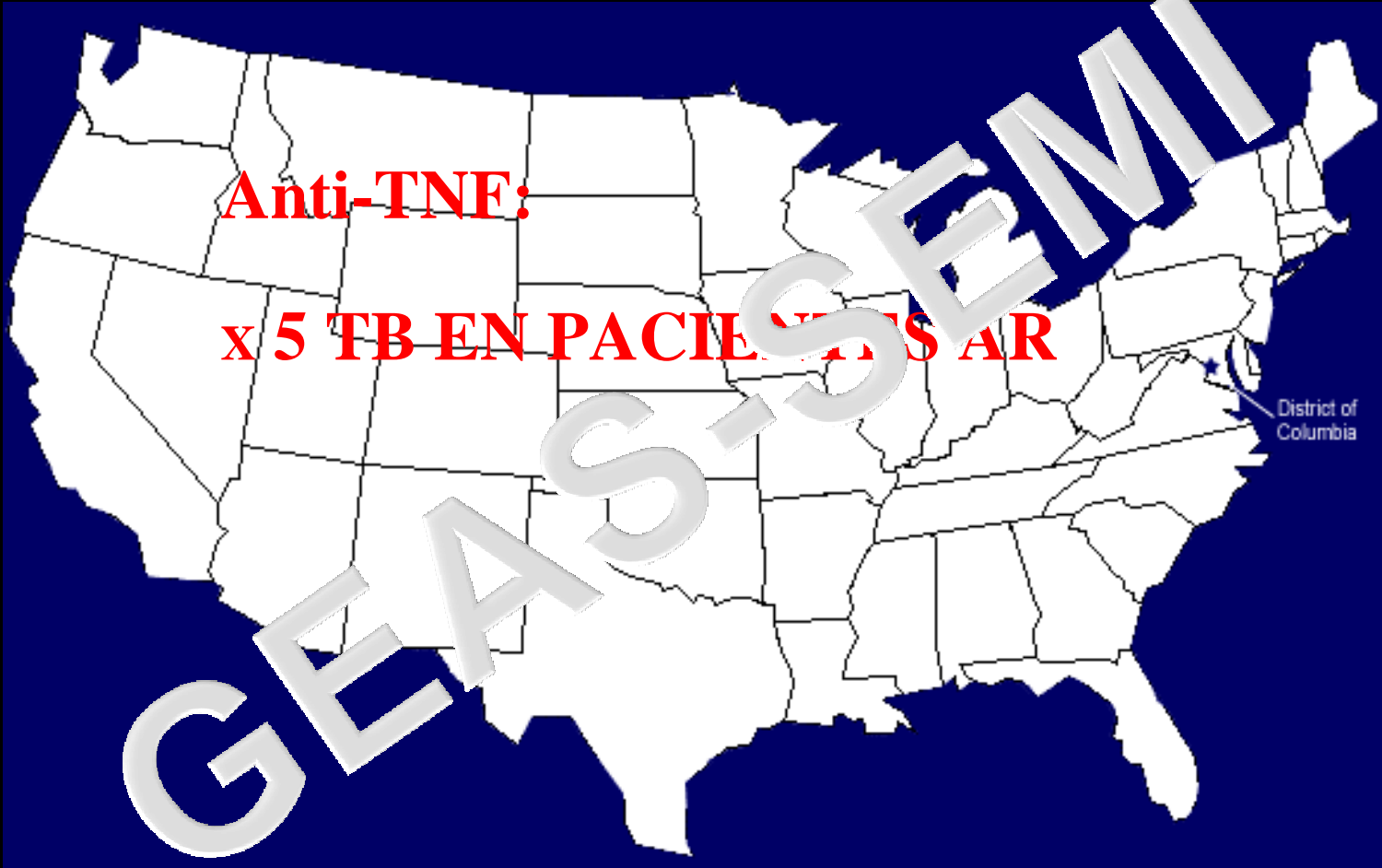
Table 2. Rate of active TB in the BIOBADASER cohort before and after the official recommendations, and risk ratio for the incidence of active TB compared with the risk in the background Spanish population and in the EMECAR patients.

	Patient-years of exposure to TNF antagonists	No. of active TB cases	Active TB rate per 100 patient-years (95% CI)	IRR versus background (95% CI)	IRR versus EMECAR (95% CI)†
All TB cases					
Pre-OR	6,126	32	5.2 (3.69–7.38)	20.9 (12.0–36.8)	–
Post-OR	1,699	2	1.17 (0.29–4.70)	4.7 (0.5–18.9)	–
IRR _{recommendations} ‡	–	–	0.22 (0.03–0.88)	–	–
TB cases with RA only					
Pre-OR	1,300	27	2.07 (1.38–3.11)	22.6 (12.6–40.6)	6.2 (2.6–16.9)
Post-OR	1,199	1	0.08 (0.01–0.76)	3.8 (0.1–23.3)	1.0 (0.02–8.2)
IRR _{recommendations} ‡	–	–	0.17 (0.004–1.02)	–	–

* TB = tuberculosis; IRR = incidence risk ratio; 95% CI = 95% confidence interval (see Table 1 for other definitions).

† EMECAR patients were patients with rheumatoid arthritis (RA) who were not treated with TNF antagonists and were followed up for 5 years in the Morbidity and Clinical Expression of Rheumatoid Arthritis study.

‡ IRR_{recommendations} = incidence risk ratio comparing the rates of active TB before (pre-OR) and following (post-OR) the official recommendations implemented on March 1, 2002 for the management of latent TB infection.



Anti-TNE:

x 5 TB EN PACIENTES AR

District of
Columbia



MMWR™

- Realizar Mantoux antes tratamiento anti-TNF.
- Mantoux > 5mm: Infección tuberculosa latente.
- Isoniacida 9 meses.
- Si Mantoux negativo: isoniacida si contexto epidemiológico o clínico sugestivo.

GEASIS-SEMINAR

Arthritis Rheum 2005; 52:2968-2074

BTS GUIDELINES

BTS recommendations for assessing risk and for managing *Mycobacterium tuberculosis* infection and disease in patients due to start anti-TNF- α treatment

British Thoracic Society Standards of Care Committee*



2010; 10: 1-8. doi: 10.1136/thx.2005.046797

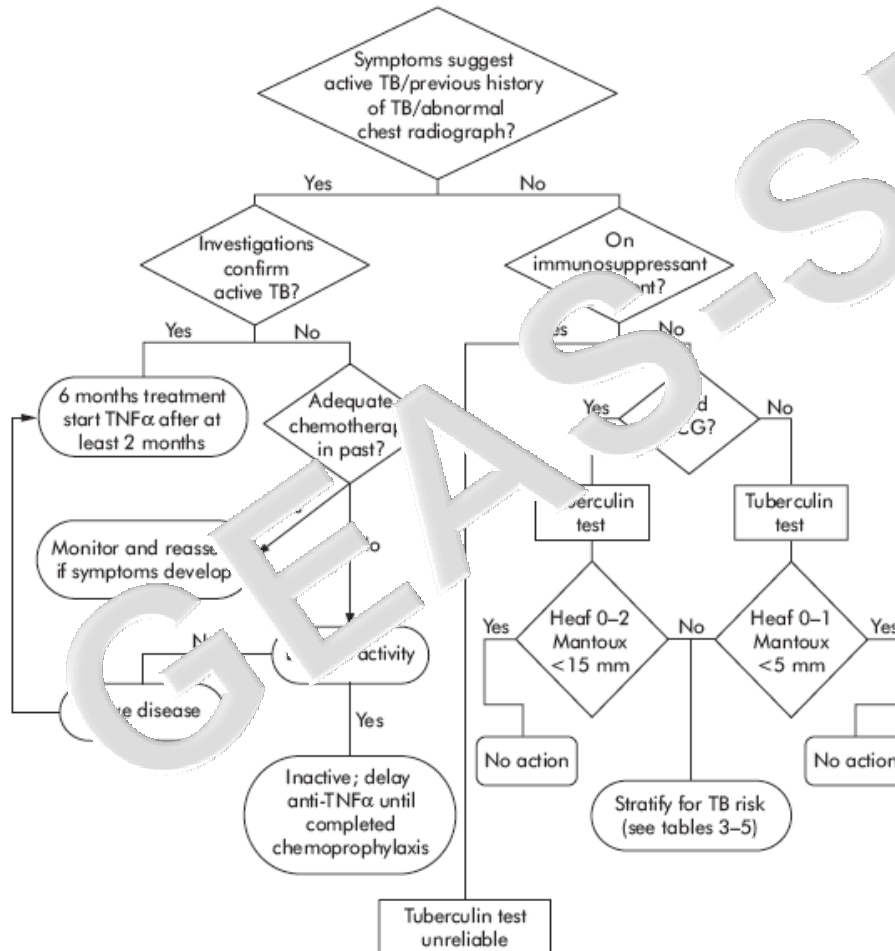


Table 5 Sample calculations based on data in tables 3 and 4

Case type	Annual risk of TB disease/100 000	TB risk adjusted $\times 5$ for anti-TNF effect	Risks of prophylaxis /100 000 (table 6)	Regimen
White Age 55–74 UK born	7	35	278	Observation
ISC Age >35 In UK 3 years	593	2965	278	Prophylaxis
Black African Age 35–54	8	40	278	Prophylaxis
Other ethnicities Age >35 in UK 3 years	195	975	278	Observation

The weighted average risk for prophylaxis with isoniazid (6H) is 278/100 000 which is used for these calculations. The weighted average risk for rifampicin/isoniazid (3RH) is higher at 1766/100 000, but this regimen may need to be considered if a shorter duration of chemoprophylaxis is needed on clinical grounds (see section 4.6).

Tuberculosis in patients receiving anti-TNF agents despite chemoprophylaxis

613 pacientes: criterios 45 profilaxis (36) Mantoux + 10 mm

Table Characteristics of patients with active TB

No.	Age, years	Sex	Collagen vascular disease	TST	CXR	Site of active TB	Chemoprophylaxis regimen	Interval to active TB, months*	Medication for collagen vascular disease
1	51	M	Ankylosing spondylitis	+	Apical fibrotic lesions	Pulmonary	INH+RMP [†]	3	MTX-CS, adalimumab
2	43	M	RA	+	Apical fibrotic lesions Pleural calcification	Pulmonary	INH+RMP [†]	4	MTX-CS, infliximab
3	50	F	Ankylosing spondylitis	-	Normal	Lymph nodes	None	12	MTX-CS, infliximab
4	62	F	Psoriatic arthritis	+	Normal	Nasopharynx	Inadequate	8	CS, infliximab
5	60	F	RA	+	Normal	Pleural effusion	INH [‡]	3	MTX, infliximab
6	52	F	RA	+	Apical fibrotic lesions	Pulmonary	INH+RMP [†]	2	MTX-CS, infliximab
7	59	F	Ankylosing spondylitis	+	Normal	Oropharynx	Inadequate	4	MTX-CS, adalimumab
8	48	M	Reiter's syndrome	+	Normal	Pulmonary	INH [‡]	2	CS, adalimumab
9	54	M	Ankylosing spondylitis	+	Normal	Pulmonary	INH [‡]	3	MTX-CS, infliximab
10	63	F	RA	+	Normal	Spleen	None	18	MTX-CS, infliximab
11	49	F	Psoriatic arthritis	+	Normal	Pulmonary	INH [‡]	35	CS, infliximab

* Interval between starting anti-TNF agent and diagnosis of active TB.

[†] 3-month regimen.

[‡] 6-month regimen.

TB = tuberculosis; RA = Rheumatoid arthritis, TST = tuberculin skin test; CXR = chest radiograph; INH = isoniazid; RMP = rifampicin; MTX = methotrexate (up to 15 mg/week), CS = corticosteroids (up to 10 mg/day prednisolone).

80 pacientes



Uso de las terapias biológicas en enfermedades autoinmunes sistémicas

Listado de efectos adversos en los pacientes incluidos en el Registro (fecha actualización: 31-12-07)

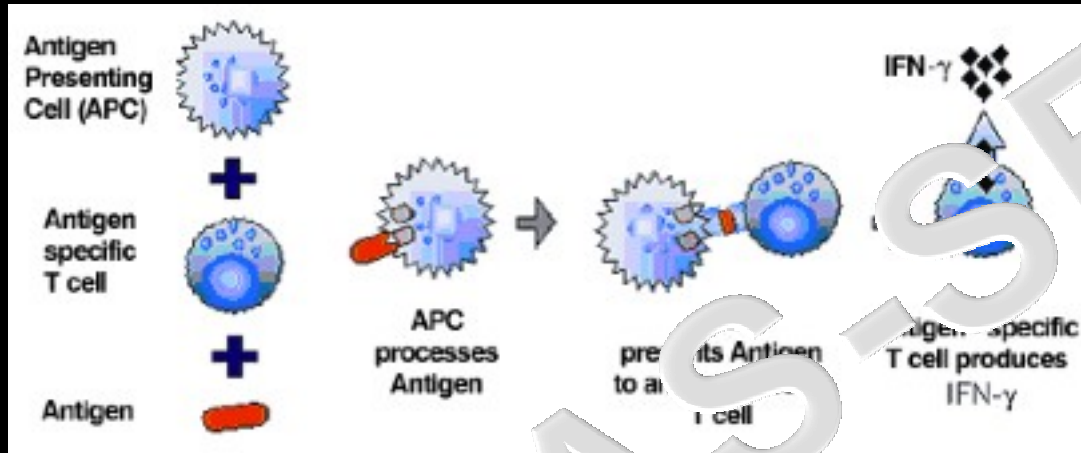
ID	EAS	BIOLOGICO	Sexo	Edad	Efecto adverso	Relación con agente
3	STILL ADULTO	infiximab	F	45	Reacc hipersensibilidad	Probable
8	PAN	infiximab	F	43	Reacc hipersensibilidad	Probable
9	VOGT-KOYANAGI	infiximab	F	37	Reacc hipersensibilidad	Probable
12	SJOGREN	rituximab	F	82	Neoplasia pulmonar	Improbable
14	DERMATOMIOSITIS	infiximab	F	63	Reacc hipersensibilidad	Probable
15	DERMATOMIOSITIS	infiximab	F	59	Infección x herpes	Probable
16	DERMATOMIOSITIS	etanercept	F	75	Colecistitis CMV	Probable
19	WEGENER	rituximab	F	59	Vaginitis x trichomona	Probable
20	LES	rituximab	F	42	Colitis	Poco probable
33	WEGENER	rituximab	M	48	Muerte súbita (sospecha TEP)	Poco probable
35	BEH	etanercept	F	24	Sospecha enf. desmielinizante	Probable
37	LES	rituximab	F	24	Neumonía intrahospitalaria	Probable
43	LES	rituximab	F	51	ITU repetición	Probable
45	LES	rituximab	F	52	Neumonía	Probable
50	LES	rituximab	F	31	Reacc hipersensibilidad	Probable
51	LES	rituximab	M	29	Fascitis necrotizante	Improbable
78	VHC	rituximab	F	66	ENdocarditis MRSA	Probable

REGISTRO IBIOGEAS

Registro Internacional de Pacientes con EAS Tratados con Agentes Biológicos

SAD	Biological agent	Infections
Horton	ETANERCEPT	4 infections (mild respir or ITU)
PAN	ETANERCEPT	Meningitis (cryptococo sp)
Polimyositis	ETANERCEPT	recurent esophageal candid
Polymyalgia R	ETANERCEPT	2 cystitis (ns) and 1 flu syndr
Sarcoidosis	ETANERCEPT	cellulitis lower extremity
Sarcoidosis	ETANERCEPT	respiratory infection
Still	ETANERCEPT	Meningitis
Wegener	ETANERCEPT	infections (4)
Sjogren	ETANERCEPT	conjunctivitis
Behçet	INFLIXIMAB	herpetic keratitis
Behçet	INFLIXIMAB	
Behçet	INFLIXIMAB	herpes zoster (1)
Behçet	INFLIXIMAB	herpes zoster
Behçet	INFLIXIMAB	respiratory infections
Behçet	INFLIXIMAB	herpetic keratitis plus otitis; 1 severe herpes; 1 recurrent ITU
Behçet	INFLIXIMAB	secondary TBC; pumonary legionella
Behçet	INFLIXIMAB	minor IRVA in 4
DM	INFLIXIMAB	osteomyelitis (MRSA + pseudomona) and pneumonia (MRSA)
DM	INFLIXIMAB	aspiration pneumonia and multi-organism bacteremia (Kb pneumoniae, S epidermidis, E coli, and S aureus)
Systemic sclerosis	INFLIXIMAB	peritonitis (candida albicans)
Behçet	INFLIXIMAB	mild infections (17), zoster keratitis (1), bronchitis (1), infected hematoma (SA) and then pleunopneumonia (1)
Behçet	INFLIXIMAB	ITU (E.coli in one, no specifi in 2)
Behçet	INFLIXIMAB	Pneumonia x 2 (H influenza) (1), recurr ITU (klebsiella) (1), leg cut abcess (MRSA) (1), endoftalmitis (nocardia) (1), broncopneumonia (ns) (1), skin ulcer (SA) + itu (E coli) (1), diarreic (ns) (1)
Policondritis	INFLIXIMAB	parasternal abcess (MRSA) and then multiple abcess and pneumonia
Polimiositis	INFLIXIMAB	pumonary (mycobacterium pellegrinum)
Sarcoidosis	INFLIXIMAB	TBC
Sarcoidosis	INFLIXIMAB	Oral candida (1)
Sarcoidosis	INFLIXIMAB	respiratory infections (31) (ns), pneumonia (6) (ns), other (17) (ns)

Diagnóstico infección latente

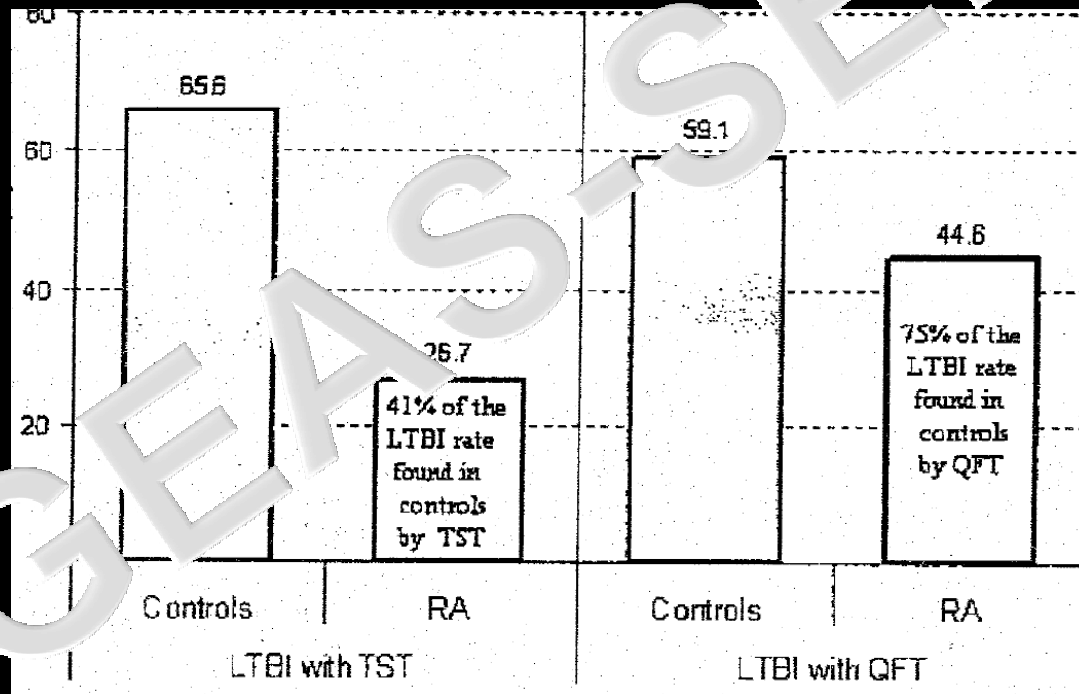


RD1:

- ESAT-6: early secretory antigen target-6
 - CFP-10: culture filtrate protein 10
 - Antígeno TB 7.7
- Quantiferon-TBGold In Tube

Comparasion of an Interferon Gamma Assay with Tuberculin Skin Testing for Detection of Tuberculosis (TB) Infection in Patients wiht Rheumatoid Arthritis in a TB-Endemic Population

J. Rheumatol 2008; 35:776



Detection of latent tuberculosis in immunosuppressed patients with autoimmune diseases: performance of a *Mycobacterium tuberculosis* antigen-specific interferon- γ assay

Ann Rheum Dis 2008;67:84-

142 patients

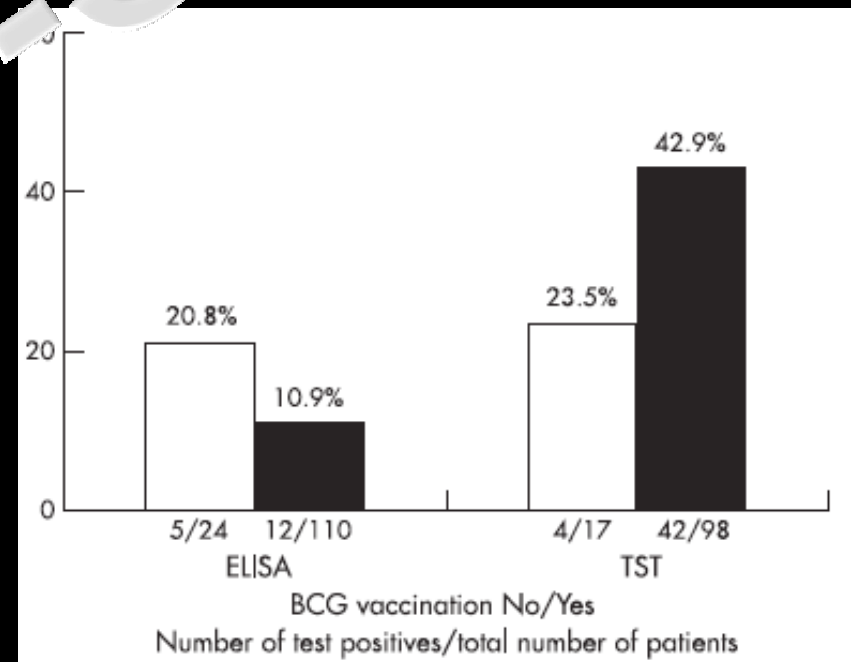
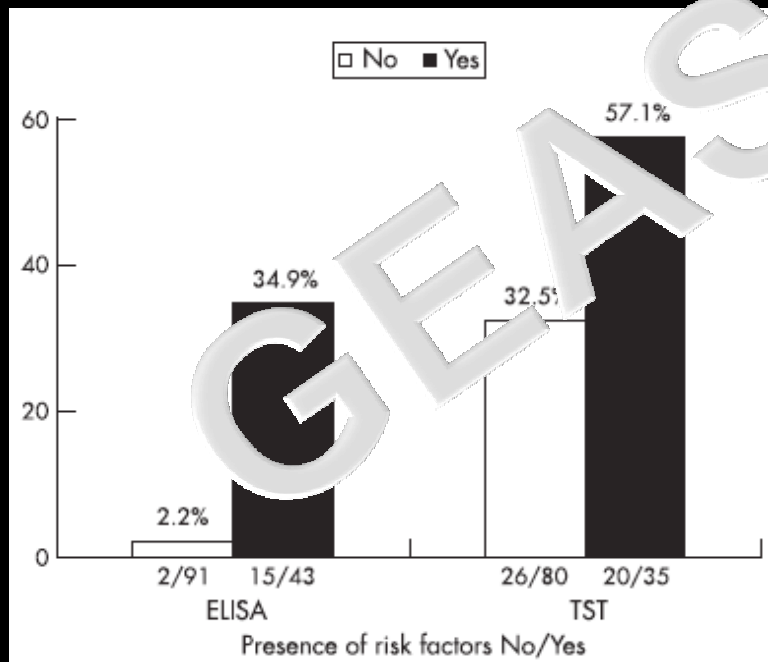


Table 2 Interferon γ assay and the tuberculin skin test according to the presence/absence of risk factors for latent *Mycobacterium tuberculosis* infection and BCG vaccination

Characteristic	Interferon γ assay (IFN γ)		Tuberculin skin test (TST)		IFN γ vs TST
	Odds ratio (95% CI)	p Value	Odds ratio (95% CI)	p Value	
<i>Univariable analysis</i>					
Presence of risk factor	23.8 (5.14 to 110)	<0.001	2.77 (1.2 to 6.27)	0.015	0.009
BCG vaccination	0.47 (0.15 to 1.47)	0.20	0.74 (0.27 to 2.01)	0.14	0.025
<i>Multivariable analysis</i>					
Presence of risk factor	29.2 (5.14 to 144)	0.001	4.81 (1.80 to 12.8)	0.002	0.041
BCG vaccination	0.79 (0.4 to 1.5)	0.37	5.83 (1.46 to 23.3)	0.013	0.20

Association between the presence of any risk factor for latent tuberculosis or BCG vaccination status and results of IFN γ assay (n = 134) and TST (n = 115). Univariable analysis and multivariable analysis including both the presence of risk factors and BCG vaccination are shown. All variables were included in the multivariable analysis. An odds ratio of 23.8 indicates, for example, that patients with a risk factor for latent tuberculosis have an odds of a positive IFN γ assay, which is 23.8 times higher than the odds of patients without a risk factor. The p value for differences in performance between IFN γ assay and TST is from matched-pairs logistic regression analysis. Refer to fig 2 for numbers of patients with positive test results according to risk factor and BCG vaccination status.

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Conclusiones

- Investigación y tratamiento infección tuberculosa latente pacientes anti-TNF.
- ¿Actitud tratamiento esteroideo España?
 - Tipo EAS.
 - Tipo inmunosupresión.
 - Edad.
- Cuantificación prometedora ¿*gold estándar*?

“El que por la mañana ha conseguido conocer la verdad,
ya puede dormir por la tarde”

Confucio

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