

AAD ANNUAL MEETING **2026**

AEDV

highlights
Denver, Colorado

27 — 31
Marzo

[A un nuevo nivel de conocimiento científico]

Una iniciativa de:



Con el patrocinio de:



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Psoriasis

La llegada de los nuevos orales

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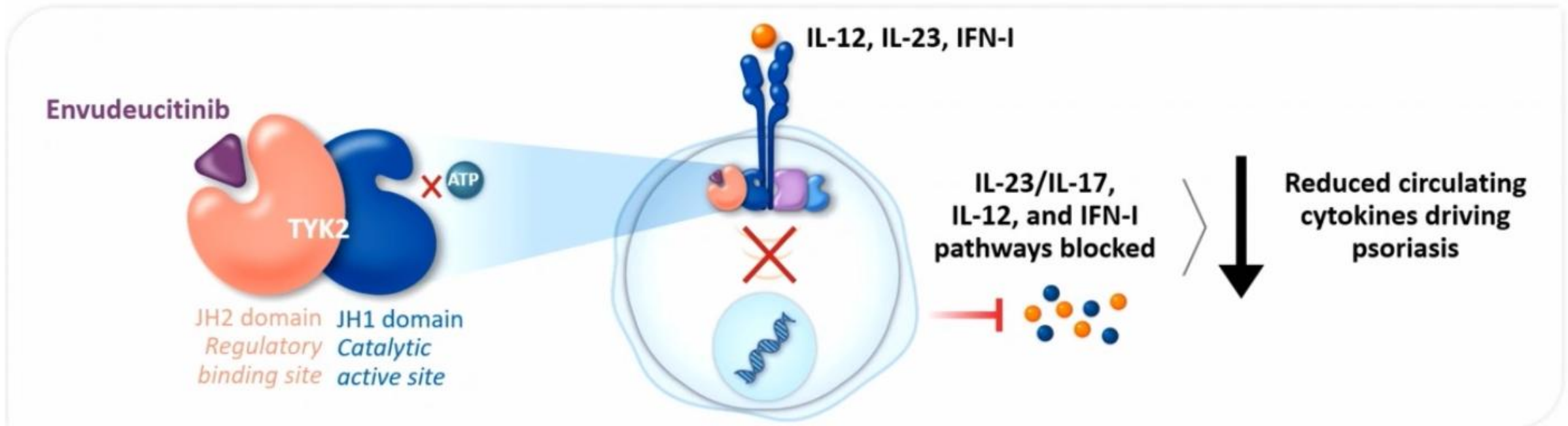
DECLARACIÓN DE CONFLICTOS DE INTERÉS

- He proporcionado asesoramiento científico/actuado como ponente/participado en ensayos clínicos de Abbvie, Almirall, Boheringuer Ingelheim, Chiesi, LeoPharma, Johnson&Johnson, KrystalBiotech, Novartis, Lilly, Loreal, Bristol Myers Squibb, UCB, Oruka Therapeutics, Takeda
- Este contenido está esponsorizado por UCB

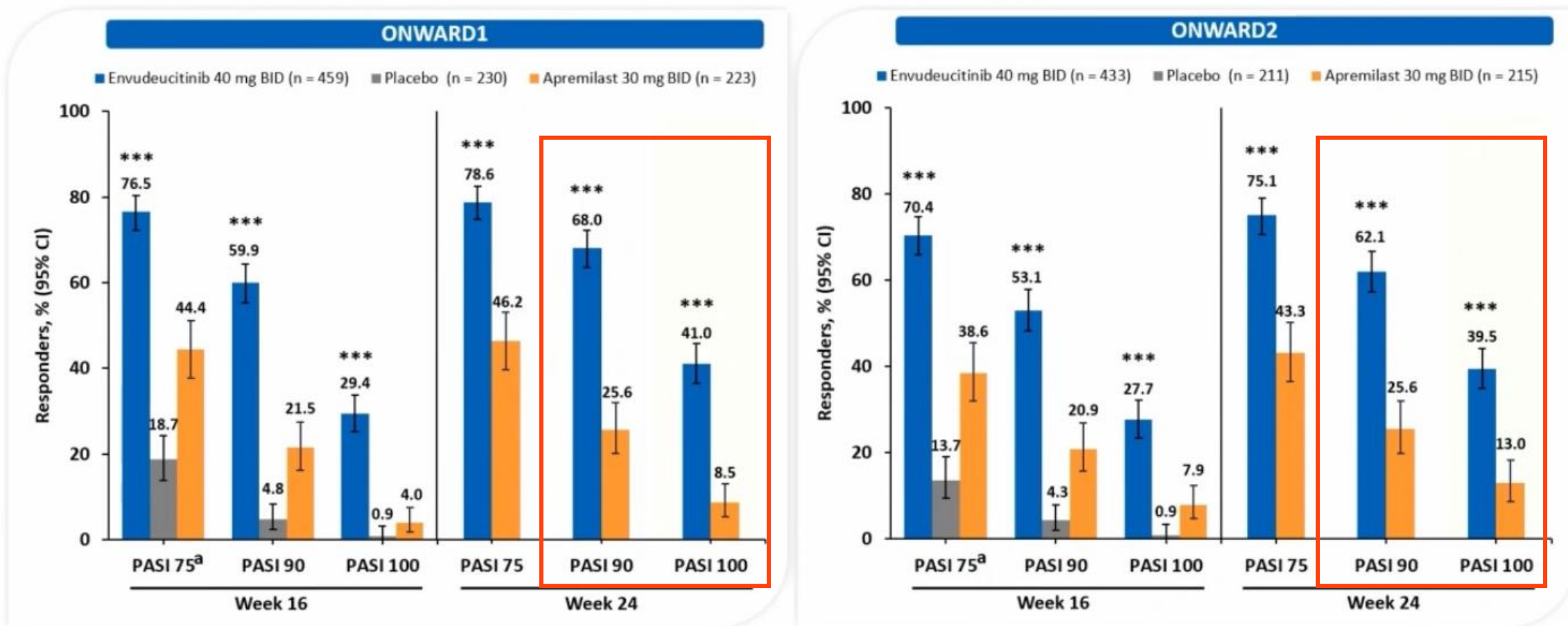
Novedades terapéuticas: inhibidores de TYK2

Envudeucitinib

- Envudeucitinib, un **inhibidor alostérico oral de TYK2** de nueva generación
- Bloqueo del eje IL-23/IL-17 pero también otras vías en menor medida



Novedades terapéuticas: inhibidores de TYK2



Intention-to-treat population. The 95% CIs and *P*-values of the treatment differences were based on the Cochran-Mantel-Haenszel test adjusted for stratification factors. Nonresponder imputation was applied for missing data. ^aCoprimary endpoint: PASI 75 at Week 16 vs placebo. ****P* < 0.0001 vs placebo and apremilast.

BID, *bis in die* (twice daily); CI, confidence interval; PASI, Psoriasis Area and Severity Index; PASI 75/90/100, ≥75%/≥90%/100% improvement in PASI.

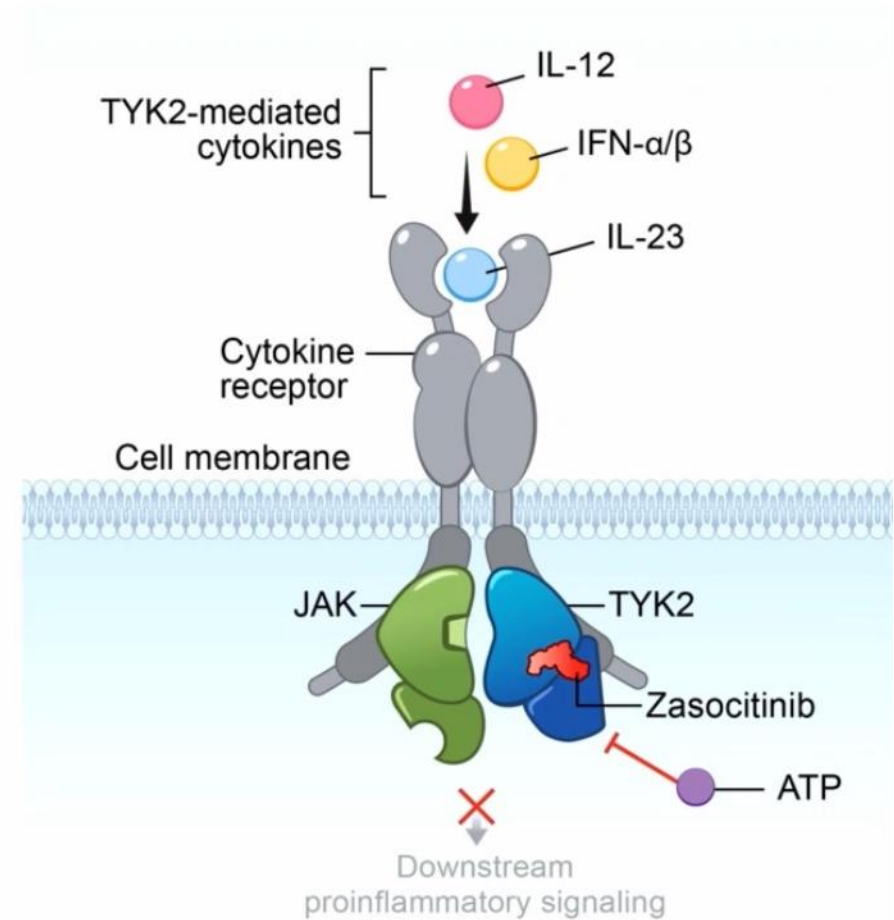
ONWARD1 and ONWARD2 Pooled Safety Through Weeks 16 and 24

n (%)	Through Week 16			Through Week 24			
	Envudeucitinib 40 mg BID n = 890	Placebo n = 441	Apremilast 30 mg BID n = 438	Envudeucitinib 40 mg BID only n = 890	Placebo to Envudeucitinib 40 mg BID n = 390	Overall Envudeucitinib 40 mg BID n = 1280	Apremilast 30 mg BID n = 438
≥1 TEAE	524 (58.9)	166 (37.6)	223 (50.9)	563 (63.3)	130 (33.3)	693 (54.1)	248 (56.6)
≥1 SAE	19 (2.1)	5 (1.1)	5 (1.1)	24 (2.7)	1 (0.3)	25 (2.0)	6 (1.4)
TEAE leading to treatment discontinuation	30 (3.4)	7 (1.6)	9 (2.1)	31 (3.5)	4 (1.0)	35 (2.7)	12 (2.7)
TEAE grade ≥3	42 (4.7)	14 (3.2)	18 (4.1)	48 (5.4)	7 (1.8)	55 (4.3)	23 (5.3)
Most-frequent TEAEs (>5%)^a							
Nasopharyngitis	64 (7.2)	21 (4.8)	16 (3.7)	92 (10.3)	18 (4.6)	110 (8.6)	26 (5.9)
Headache	92 (10.3)	11 (2.5)	40 (9.1)	97 (10.9)	11 (2.8)	108 (8.4)	42 (9.6)
Upper respiratory tract infection	43 (4.8)	7 (1.6)	16 (3.7)	57 (6.4)	2 (0.5)	59 (4.6)	21 (4.8)
Acne	53 (6.0)	3 (0.7)	3 (0.7)	60 (6.7)	17 (4.4)	77 (6.0)	3 (0.7)
Nausea	20 (2.2)	4 (0.9)	23 (5.3)	20 (2.2)	0	20 (1.6)	23 (5.3)
Diarrhea	14 (1.6)	11 (2.5)	36 (8.2)	16 (1.8)	1 (0.3)	17 (1.3)	36 (8.2)

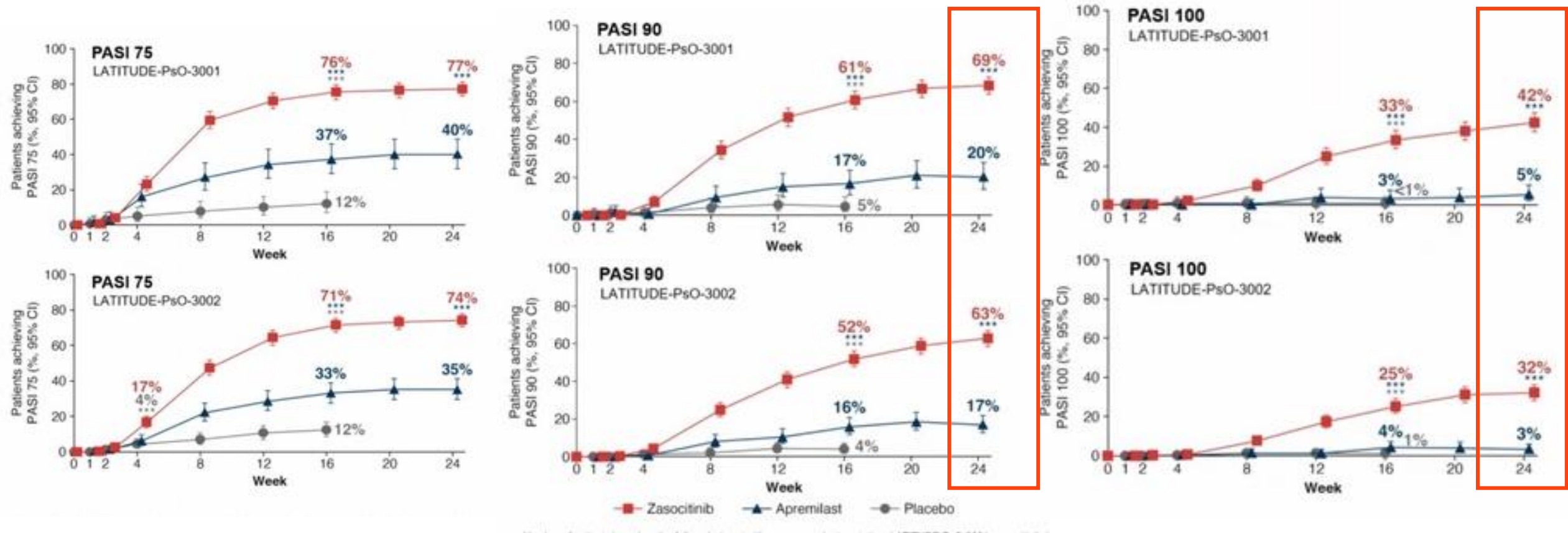
- › Envudeucitinib showed low rates of SAEs and AEs leading to discontinuation, with no clusters of events
 - No deaths; no MACE or cytopenia signals; no TB reactivation^b
- › No clinically significant laboratory abnormalities were observed across lipid, hematologic and chemistry panels, with comparable variability across treatment arms throughout the study
- › At Week 24, low incidence of serious infections (0.7%) and malignancies (0.2%) observed in patients treated with envudeucitinib

Novedades terapéuticas: zasocitinib

- Más de un millón de veces mayor selectividad de unión por TYK2 frente a JAK1, JAK2 y JAK3.
- Mantiene la inhibición durante 24 horas
- LATITUDE ensayos fase 3 vs placebo y vs apremilast en adultos con psoriasis en placas moderada-grave



Novedades terapéuticas: zasocitinib

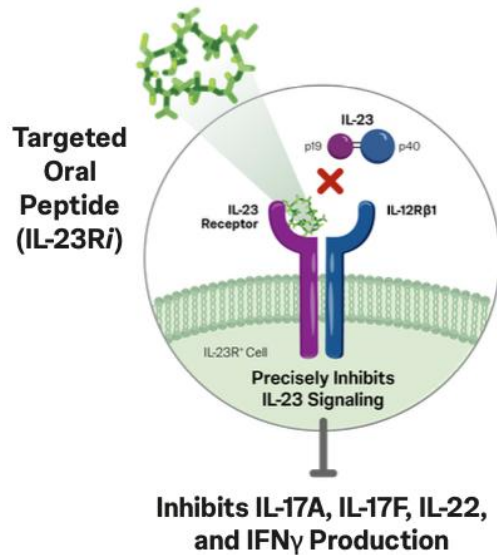


Novedades terapéuticas: zasocitinib

TEAEs from LATITUDE-PSO-3001 and 3002*	Day 0 to Week 16						Day 0 to Week 24			
	Zasocitinib (n = 970)		Apremilast (n = 412)		Placebo (n = 417)		Zasocitinib (n = 970)		Apremilast (n = 412)	
	n	% ^b (95% CI) ^c	n	% ^b (95% CI) ^c	n	% ^b (95% CI) ^c	n	% ^b (95% CI) ^c	n	% ^b (95% CI) ^c
Any TEAE	605	62.1 (59.0–65.1)	207	50.5 (45.7–55.4)	196	46.9 (42.0–51.7)	674	69.3 (66.4–72.2)	232	56.5 (51.7–61.4)
Leading to discontinuation	31	3.2 (2.1–4.3)	11	2.6 (1.1–4.2)	3	< 1 (0.0–1.6)	36	3.7 (2.5–4.9)	13	1.3 (1.4–4.7)
SAE	29	3.0 (1.9–4.1)	6	1.5 (0.3–2.7)	2	< 1 (0.1–1.7)	35	3.6 (2.4–4.8)	7	1.7 (0.5–3.0)
Death	1 ^d	< 1 (0.0–0.6) ^d	0	0 (0.0–0.9)	0	0 (0.0–0.9)	1 ^d	< 1 (0.0–0.6) ^d	0	0 (0.0–0.9)
Most frequent TEAE (≥ 5%)^a										
URTI	100	10.1 (8.2–12.0)	24	6.0 (3.7–8.3)	13	3.2 (1.5–4.8)	123	12.5 (10.4–14.6)	29	7.4 (4.8–10.0)
Acne	62	6.5 (5.0–8.1)	3	< 1 (0.0–1.7)	1	< 1 (0.0–1.3)	70	7.3 (5.6–8.9)	3	< 1 (0.0–1.7)
Nasopharyngitis	60	6.2 (4.7–7.7)	23	5.4 (3.2–7.5)	20	4.7 (2.7–6.6)	80	8.3 (6.5–10.0)	34	7.9 (5.4–10.5)
Diarrhea	30	3.1 (2.0–4.2)	33	8.2 (5.5–10.9)	8	1.8 (0.6–3.1)	36	3.7 (2.5–4.9)	33	8.2 (5.5–10.9)
Headache	27	2.8 (1.8–3.9)	26	6.3 (4.0–8.7)	8	1.9 (0.6–3.2)	32	3.3 (2.2–4.5)	28	6.8 (4.4–9.3)
Nausea	20	2.1 (1.2–3.0)	23	5.5 (3.3–7.8)	5	1.2 (0.1–2.2)	23	2.4 (1.4–3.4)	24	5.8 (3.5–8.1)

Icotrokinra en adultos a 52 semanas

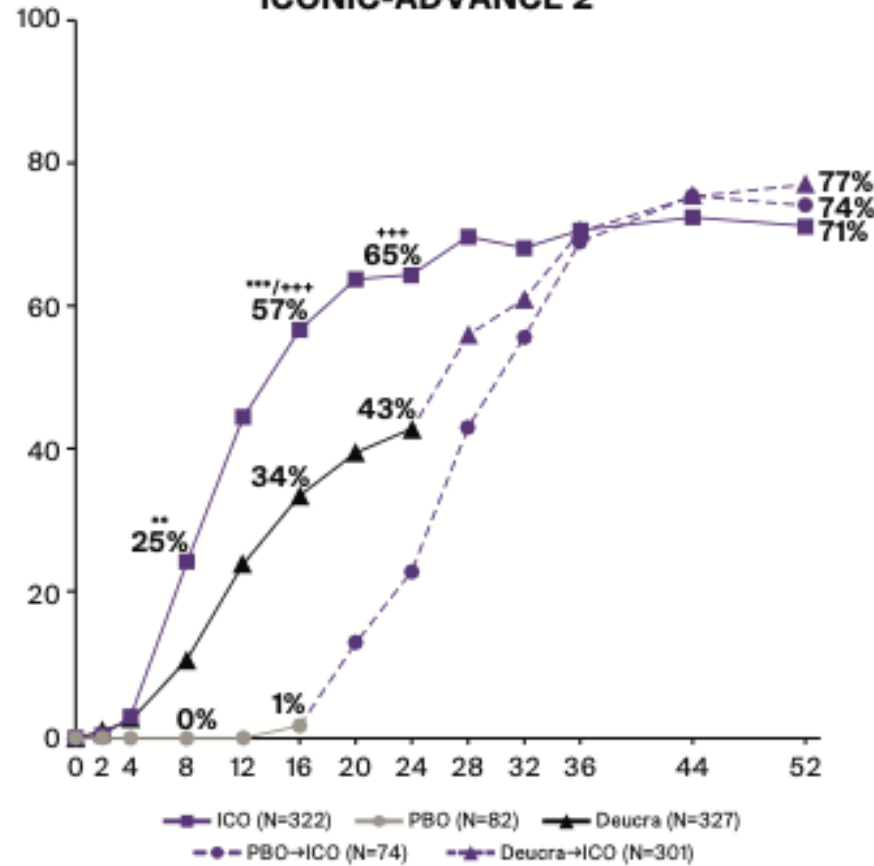
Icotrokinra Blocks IL-23 From Binding to its Receptor



IFN=interferon, *IL-12Rβ1*=interleukin-12 receptor beta 1, *IL-23Ri*=interleukin-23 receptor inhibitor.

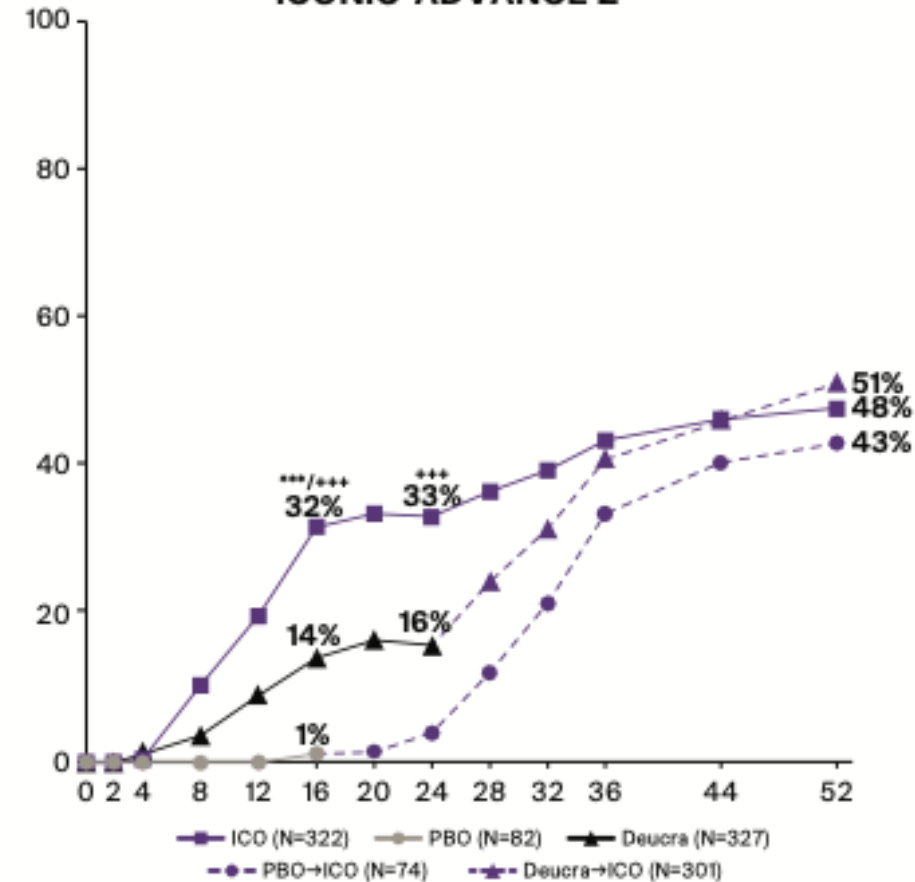
PASI 90

ICONIC-ADVANCE 2

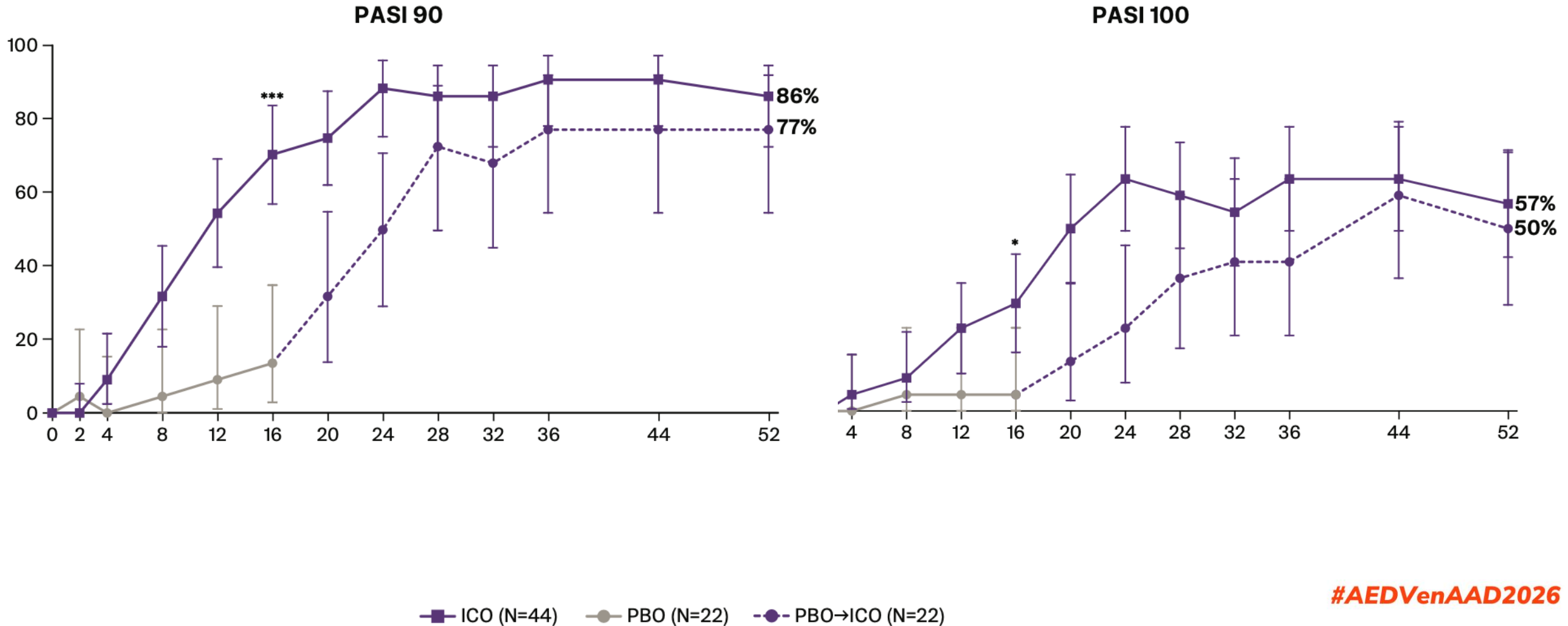


PASI 100

ICONIC-ADVANCE 2



Icotrokinra en adolescentes > 12 años

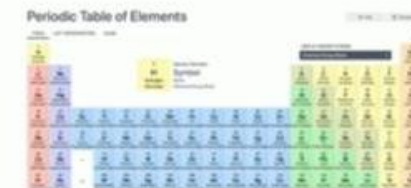


The ICO AE profile in adolescents was similar to PBO through W16, and consistent between W16 & W52

AEs Through W52: Adolescents ^a	PBO-Controlled (Through W16)		Through W52
	PBO (N=22)	ICO (N=44)	ICO Combined (N=66) ^a
Mean weeks / total PY of follow-up	16.2 / 6.8	16.2 / 13.7	46.3 / 58.6
Any AE	16 (73%)	22 (50%)	46 (70%)
Incidence/100 PY (95% CI) ^{b,c}	521 (266, 776)	238 (139, 338)	164 (117, 212)
Serious AE	0	2 (5%)	4 (6%)
Incidence/100 PY (95% CI) ^{b,d}	0 (0, 44)	15 (2, 54)	7 (2, 18)
AE leading to discontinuation	0	0	0
Incidence/100 PY (95% CI) ^{b,d}	0 (0, 44)	0 (0, 22)	0 (0, 5)
Infection	6 (27%)	14 (32%)	31 (47%)
Incidence/100 PY (95% CI) ^{b,c}	116 (23, 209)	130 (62, 198)	78 (50, 105)
Serious infection	0	0	0
Incidence/100 PY (95% CI) ^{b,d}	0 (0, 44)	0 (0, 22)	0 (0, 5)
Gastrointestinal AE	1 (5%)	2 (5%)	5 (8%)
Incidence/100 PY (95% CI) ^{b,d}	15 (<1, 85)	15 (2, 53)	9 (3, 21)
Malignancy	0	0	0
Incidence/100 PY (95% CI) ^{b,d}	0 (0, 44)	0 (0, 22)	0 (0, 5)

Data shown are n (%), unless otherwise noted. Safety analysis set included all randomized and treated pts. ^aIncludes pts receiving ICO through W52 and data after W16 for pts receiving PBO who transitioned to ICO. ^bIncidence/100 PY: (number of pts with AEs/total PY at risk) × 100. ^cCI's were based on a Wald statistic using the normal assumption. ^dCI's were based on an exact method assuming that the observed number of events follows a Poisson distribution.

What therapies can work for PPP?



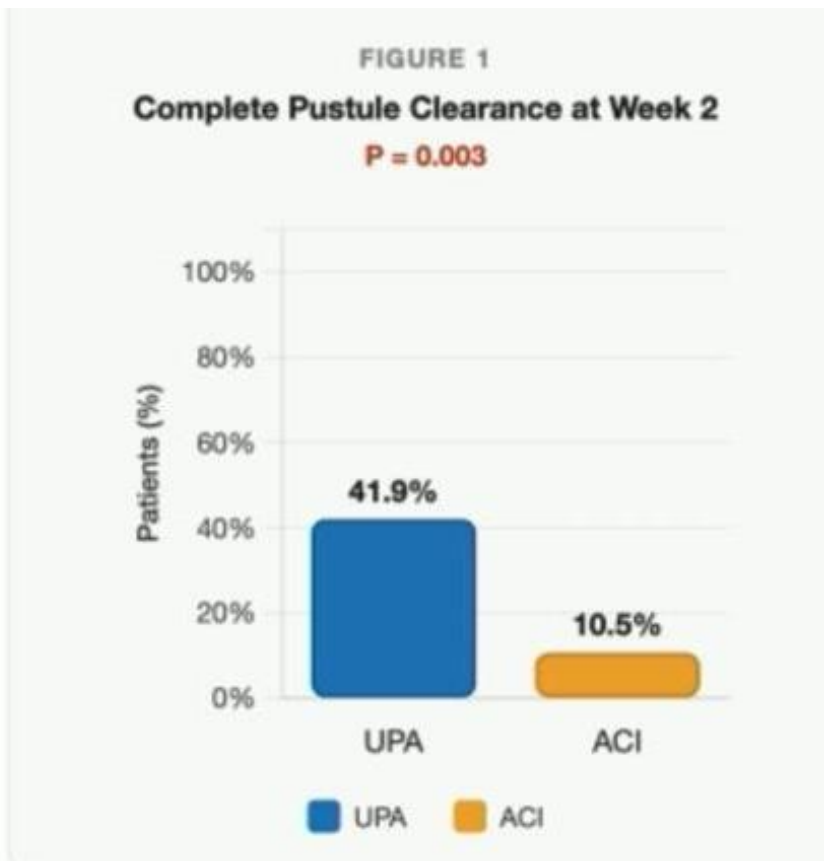
There's no green light here...

topicals

- Cs**
broadband-ultraviolet-b
- D**
vitamin D analog
- A**
vitamin A analog
- Cs-D**
Corticosteroid-vitamin D
- Cs-A**
Corticosteroid-vitamin A
- Rf**
Roflumilast
PDE-4 inhibitor
- Tf**
tapinarof
Aryl hydrocarbon receptor (AhR) agonist

1Pu Psoralen + Ultraviolet-A	2Bb broadband-ultraviolet-b	3Nb narrowband ultraviolet-b	4Exc excimer 308nm ultraviolet-b	5Hh handheld/home ultraviolet-b	
6Mtx methotrexate antimetabolite	7Aci acitretin retinoid	8Cya cyclosporine calcineurin inhibitor	16Apr apremilast phosphodiesterase-4 inhibitor	22Deu Deucravacitinib TYK2 inhibitor	23Upa Upadacitinib JAK inhibitor (AD, PsA)
9Etn etanercept TNF-a inhibitor	10Ada adalimumab TNF-a inhibitor	11Cer certolizumab pegol TNF-a inhibitor	12Inf infliximab TNF-a inhibitor	13Gol golimumab TNF-a inhibitor	
14Ust ustekinumab interleukin 12/23 inhibitor	21Til tildrakizumab interleukin-23 inhibitor	18Gus guselkumab interleukin-23 inhibitor	20Ris risankizumab interleukin-23 inhibitor	26lco Icotrokinra Oral interleukin-23 receptor antagonist	
15Sec Secukinumab interleukin-17 inhibitor	17Ixe Ixekizumab interleukin-17 inhibitor	19Bro brodalumab interleukin-17 receptor inhibitor	25Bim bimekizumab interleukin-17 inhibitor		
24Spe Spesolimab IL36RA antagonist	Ana Anakinra IL1 antagonist				

Uso de iJAK en PPP



- Zheng Y, et al. Clinical efficacy and safety of upadacitinib in the treatment of palmoplantar pustulosis: A single-center retrospective study. *Indian J Dermatol Venereol Leprol.* 2024. PMID: 39635802.
- Huang D, Jiang X, Yang N, Wang Y, Li Y, Yi X, Guo C, Gao Y, Shi Y. Upadacitinib Versus Acitretin for the Resolution of Pustules in Palmoplantar Pustulosis During Acute Phase: A Single-Center, Open-Label Prospective Cohort Study. *Am J Clin Dermatol.* 2025 Sep;26(5):843-850. doi: 10.1007/s40257-025-00971-7. Epub 2025 Jul 23. PMID: 40702395.
- Rahbar Kooybaran N, et al. Response of palmoplantar pustulosis to upadacitinib: A case series of five patients. *J Dtsch Dermatol Ges.* 2023 Nov;21(11):1387-1392. doi: 10.1111/ddg.15176. Epub 2023 Aug 21. PMID: 37605445.
- Xu Q, et al. Refractory Palmoplantar Pustulosis Successfully Treated with JAK Inhibitor Tofacitinib: A Case Series. *Infect Drug Resist.* 2023 Aug 9;16:5165-5172. PMID: 37581169

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Comorbilidades

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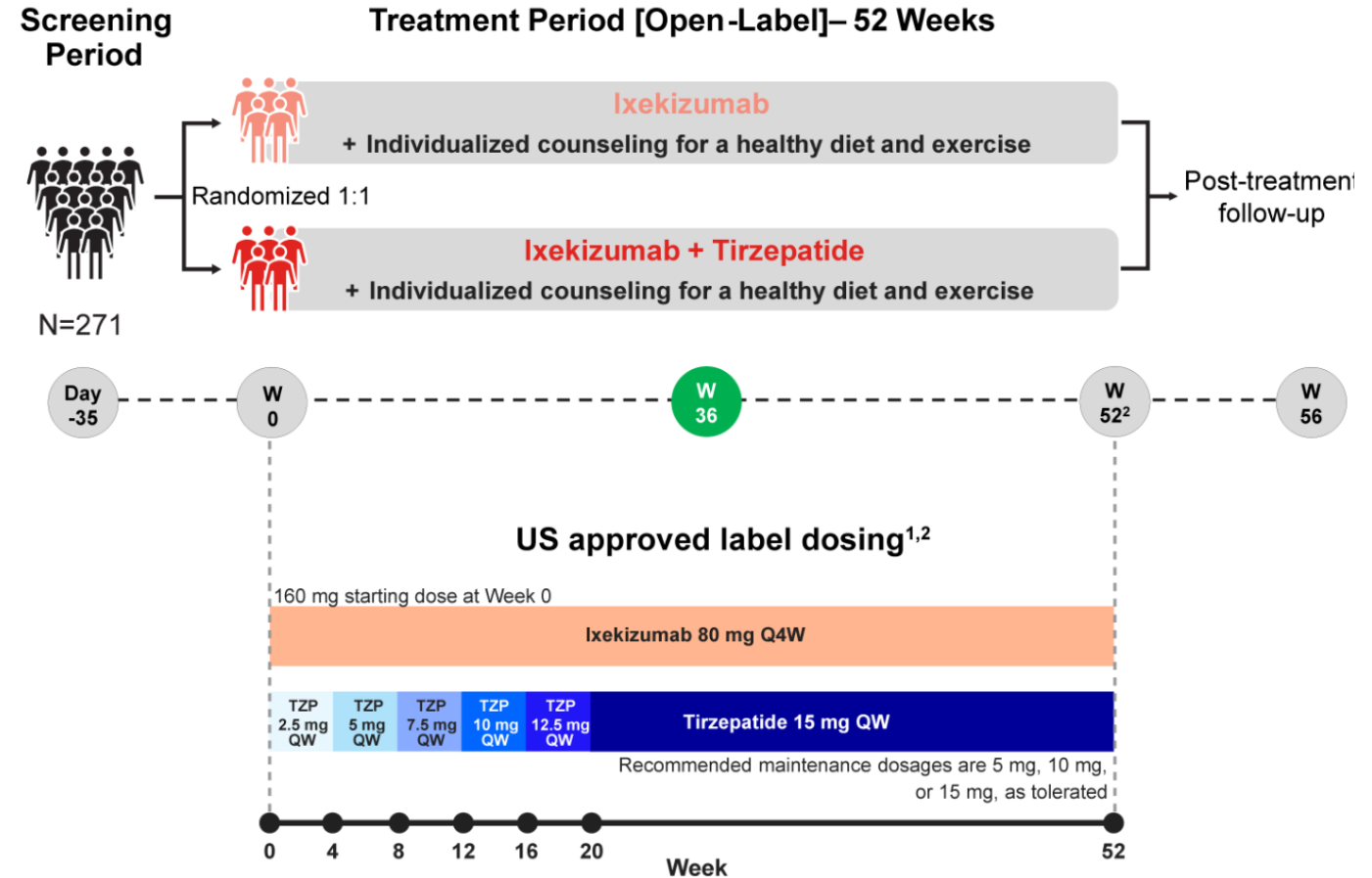
Tratamiento combinado: PsA y obesidad

El sobrepeso y la obesidad son comorbilidades frecuentes en PsO y PsA

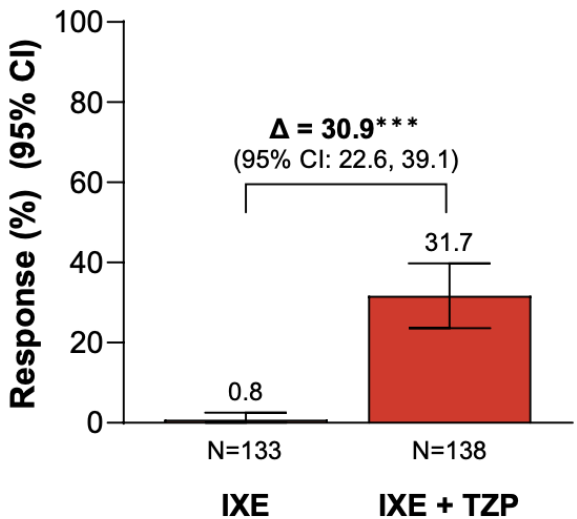
La obesidad se asocia a peor respuesta y a eventos cardiovasculares

TOGETHER PsO / PsA

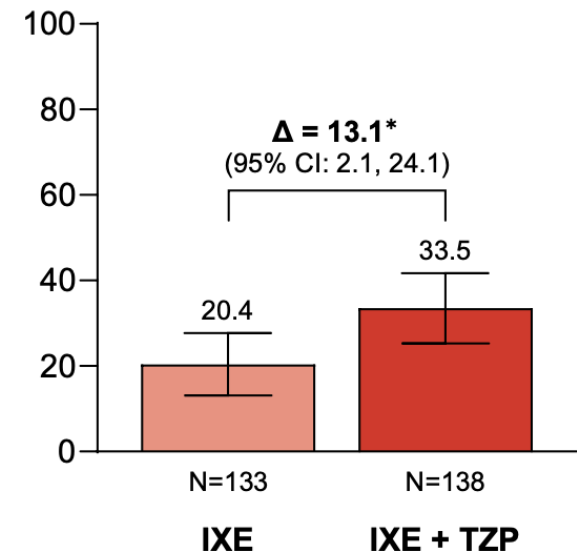
Estudia el impacto de ixekizumab vs ixekizumab + tirzepatida



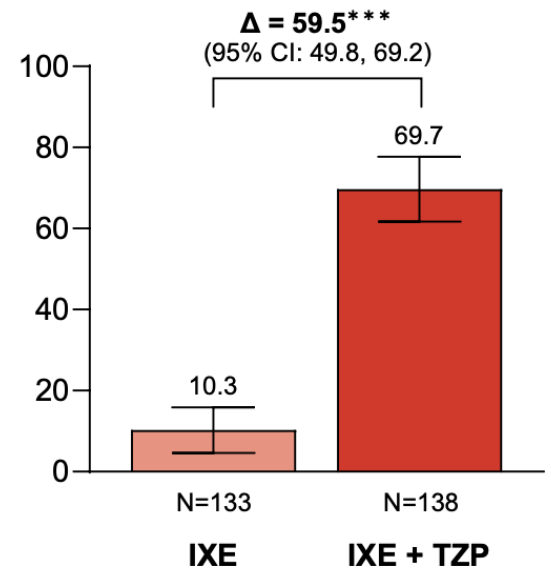
**Simultaneous⁺
ACR50 and ≥10% Weight Reduction**



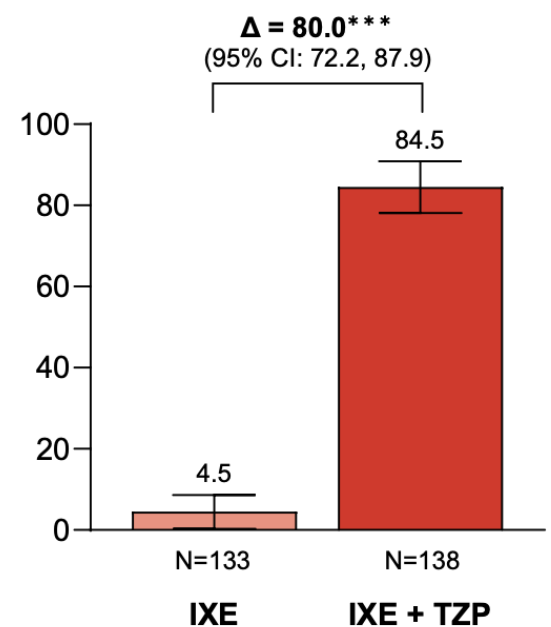
ACR50



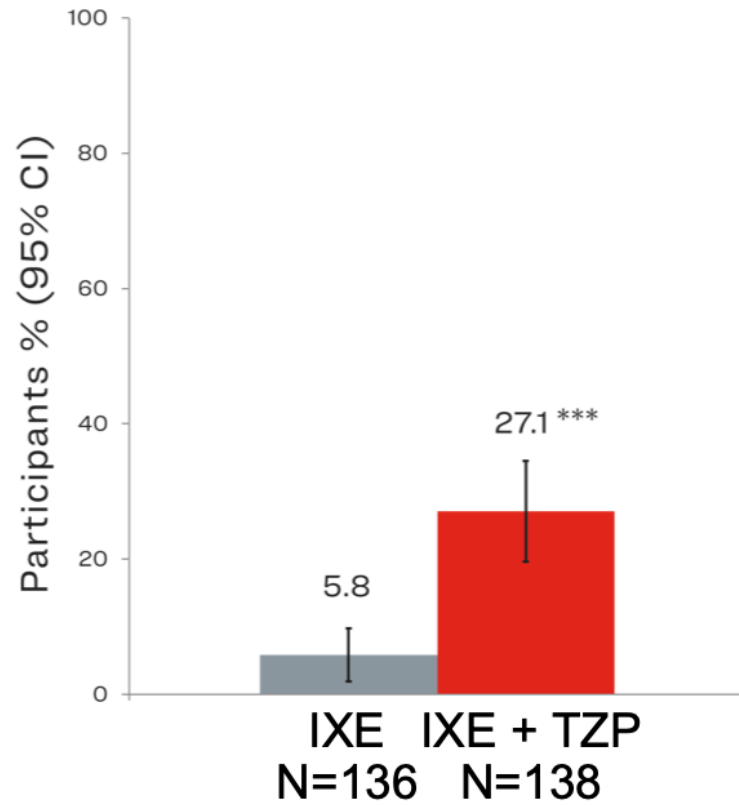
**Simultaneous
ACR20 and ≥5% Weight Reduction**



≥10% Weight Reduction

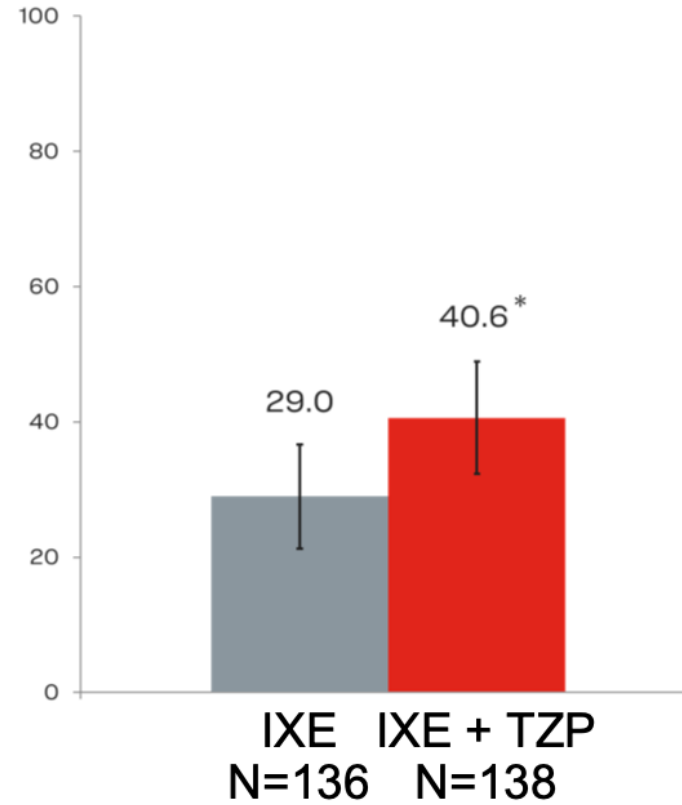


Simultaneous PASI 100 and ≥10% Weight Reduction



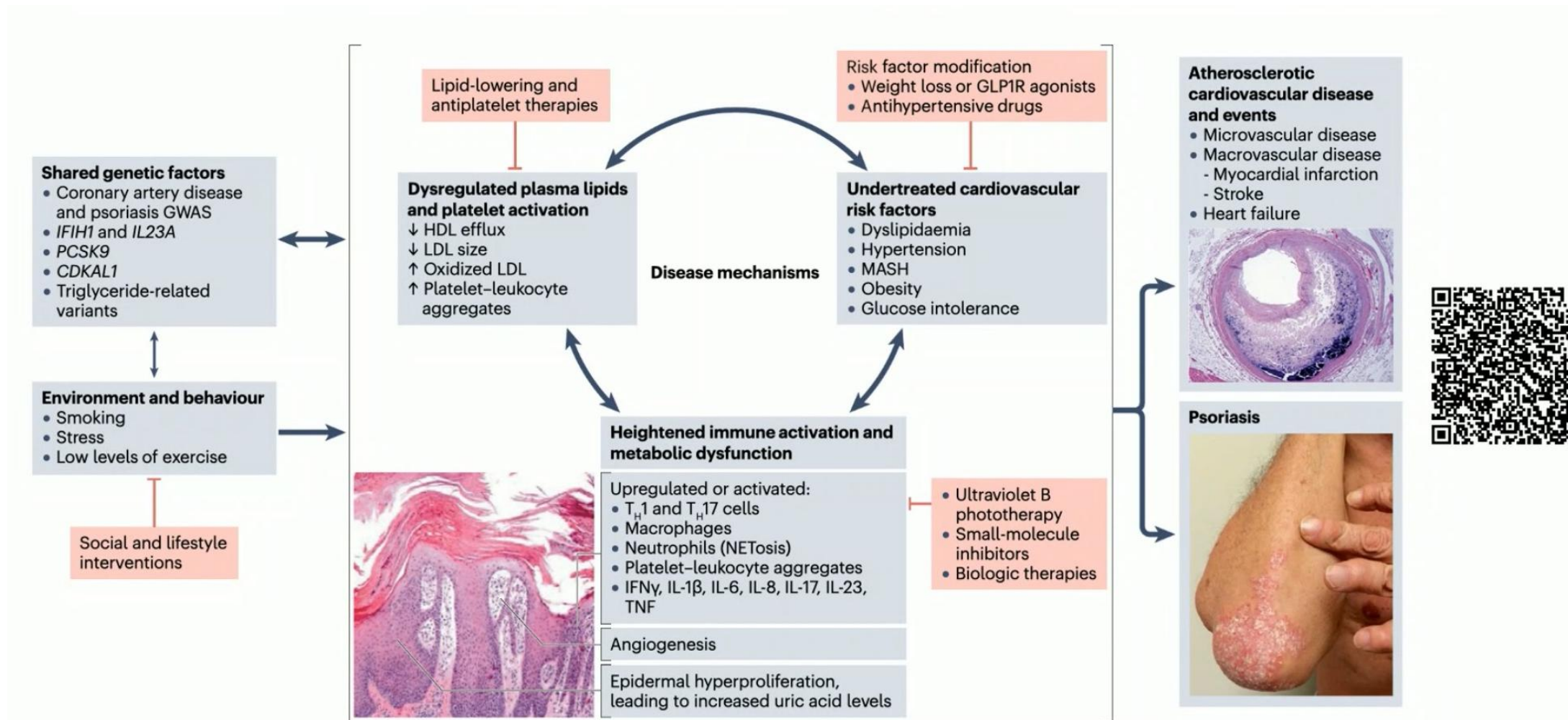
Primary Endpoint

PASI 100

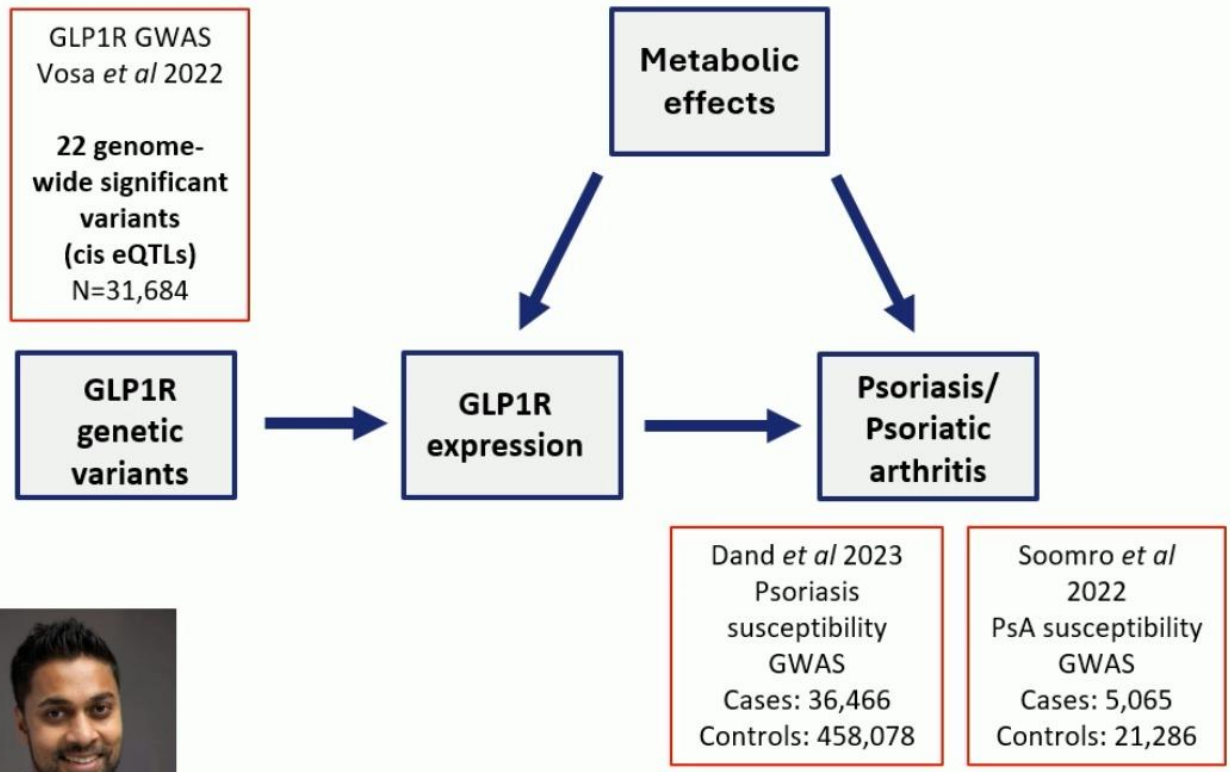


Key Secondary Endpoint

Riesgo cardiovascular



Genetically proxied GLP1 receptor signaling is protective of psoriasis & psoriatic arthritis but not selected inflammatory diseases

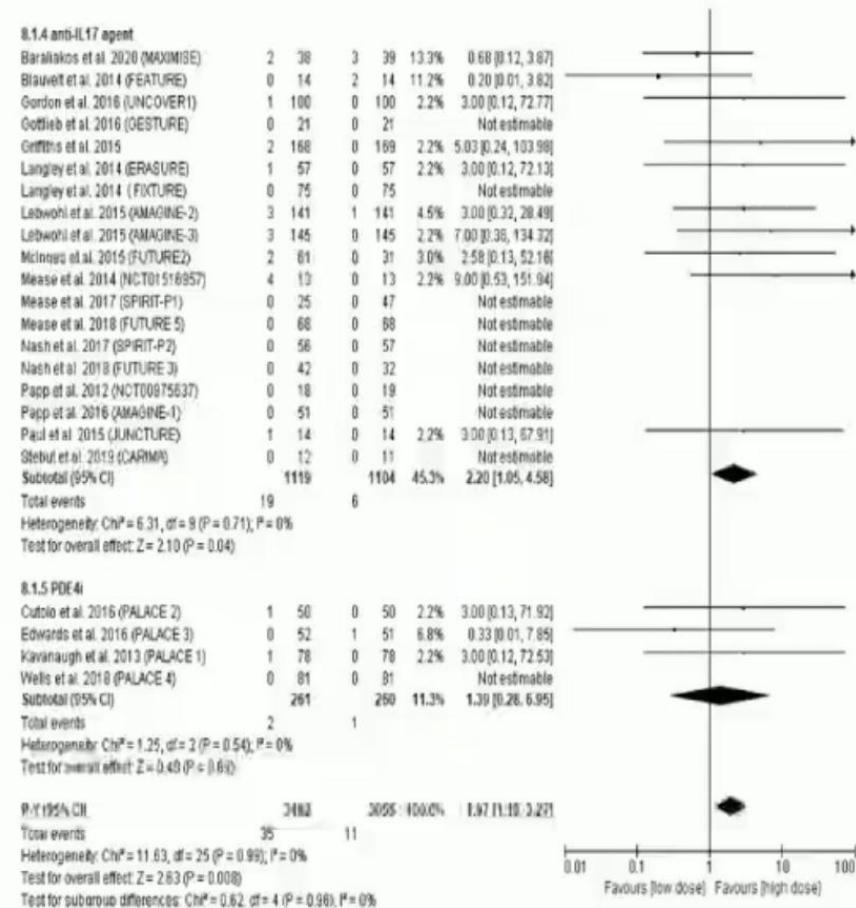


Outcome	OR (95% CI)	P-value
Psoriasis	0.72 (0.68-0.77)	1.08×10⁻²²
Psoriatic arthritis	0.48 (0.40-0.58)	5.33×10⁻¹⁵
Acne	1.04 (0.96–1.13)	0.379
Asthma	1.03 (0.99–1.07)	0.187
Atopic dermatitis	1.01 (0.96–1.05)	0.835
Crohn's disease	1.24 (1.11–1.40)	2.35×10⁻⁴
Multiple sclerosis	0.89 (0.80–1.00)	0.056
Rheumatoid arthritis	1.50 (1.35–1.67)	1.36×10⁻¹³
Ulcerative colitis	1.01 (0.96–1.05)	0.835

Meta-analysis RCTs of targeted therapies and CV events in psoriasis or PsA: No overall benefit

- 88 RCTs 10,521 person years of psoriasis or psoriatic arthritis evaluating all CV events*
- TNFi RR = 1.26, 95% CI 0.66-2.41
- JAKi RR = 1.07, 95% CI 0.37-3.11
- anti-IL-12/23 RR = 0.92, 95% CI 0.48-1.74
- anti-IL-17 RR = 1.13, 95% CI 0.62-2.07
- PDE4i RR = 0.71, 95% CI 0.23-2.18
- Results confirmed in a 2025 meta-analysis of psoriasis only RCTs

*angina pectoris, MI, congestive heart failure, carotid artery disease, aortic aneurysm, stroke, transient ischemic attack, cardiovascular death, and arrhythmia.



Artritis psoriásica

**Debate #2: Which is better at preventing
psoriatic arthritis?**



IL-23s or IL-17s

Limitaciones de los enfoques observacionales actuales

- Medición incompleta de variables de confusión, incluidas la BSA, la obesidad y la depresión
- Sesgo de selección
 - **Protopático:** el fármaco se prescribe para una manifestación precoz del desenlace que aún no ha sido detectada
 - **Confusión por indicación:** la indicación del tratamiento influye en el riesgo del desenlace
- **Sesgo colisionador:** una exposición y un desenlace influyen cada uno en una tercera variable común, y esa variable, o colisionador, se controla mediante el diseño del estudio o en el análisis (p. ej., la paradoja del tabaquismo en la APs)
- Sesgo de observación
- A menudo no hay estratificación según el mecanismo de acción del biológico

Conclusión: Algunas preguntas no se adaptan bien a los métodos observacionales. ¡Necesitamos ECA!

PROGRESSION: PSORIASIS TO PSORIATIC ARTHRITIS

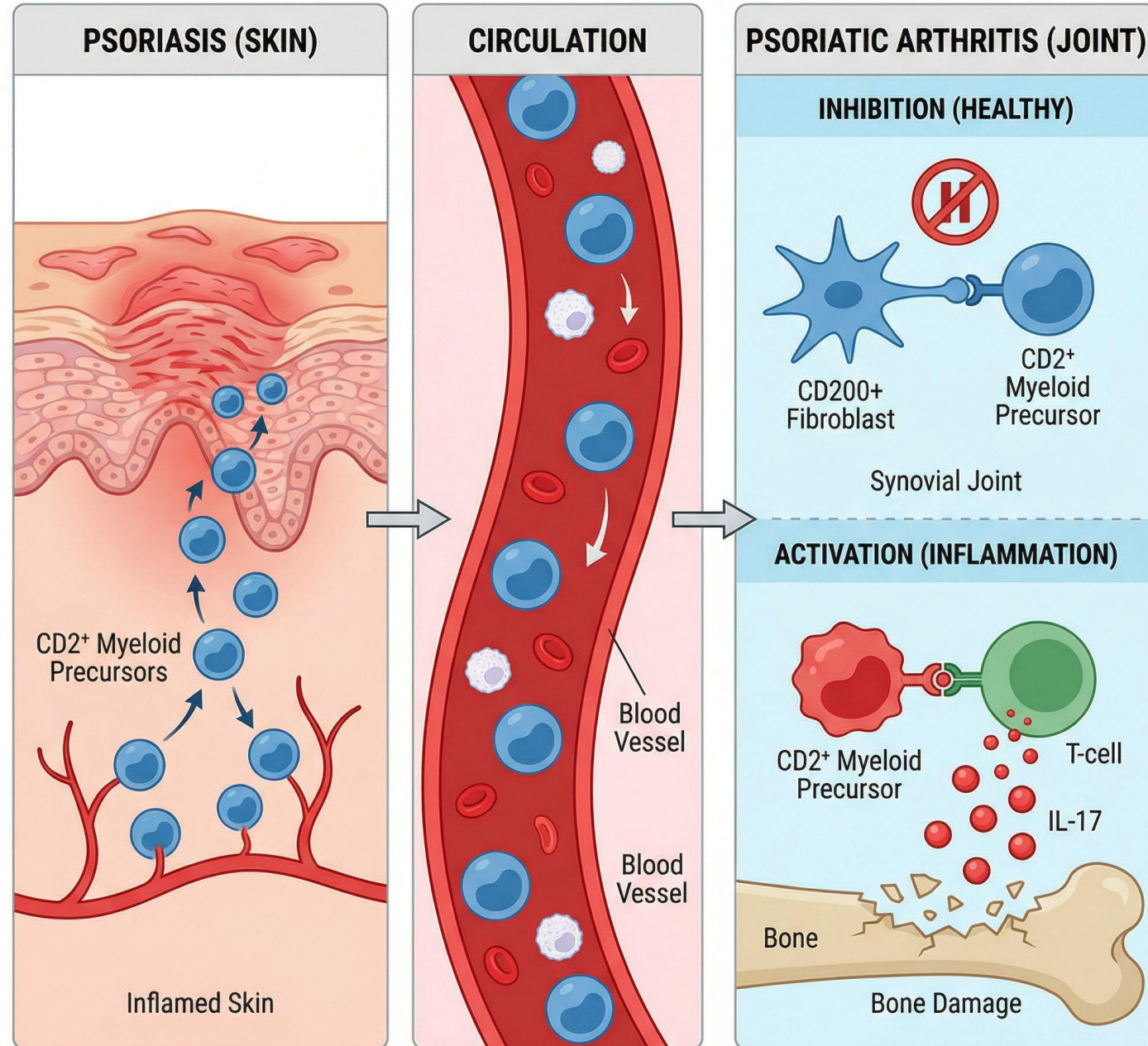
Skin-derived myeloid precursors and joint-resident fibroblasts spread psoriatic disease from skin to joints

nature immunology

Precursores precursores mieloides $CD2^+MHC$
– II^+CCR2^+ se originan en la piel psoriásica

Que haya artritis o no depende de las células locales de la articulación. En personas sanas, los **fibroblastos** $CD200^+$ frenan a estas células invasoras. Sin embargo, en quienes desarrollan artritis, estos fibroblastos protectores desaparecen.

El daño final: Al no tener freno, los precursores mieloides se activan y ordenan a los linfocitos T que produzcan grandes cantidades de **IL-17**, la molécula que finalmente destruye el hueso y el cartílago.



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Pitiriasis rubra pilaris

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Deucravacitinib tiriasis rubra pilaris

Ensayo clínico abierto, fase 2

Adultos con PRP con confirmación histológica

Deucravacitinib 6mg c/12h

Endpoint a las 24 semanas

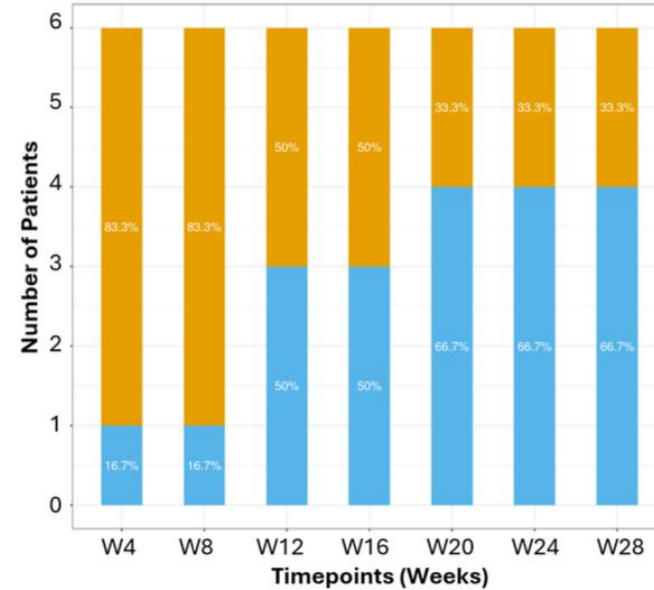
Medidas. PGA, PASI, BSA, DLQI, NRS-Itch, Skindex-16

Patients Enrolled	7
Patients Completed Study	6
Age (mean (SD)), years	59.0 (9.5)
Sex	
Female	1 (16.7%)
Male	5 (83.3%)
Race	
White (W)	5 (83.3%)
Black or African American (B)	1 (16.7%)

RESULTS



D. PGA Percentage by Timepoints



Primary Endpoint

Secondary Endpoints

	Baseline*(n=6) Mean (SD)	Week 24 (n=6) Mean (SD)	Difference (SD)	P-value
PGA	4.7 (0.8)	2.7 (1.9)	-2.0 (1.9)	0.098 ²
PASI	35.3 (11.1)	13.2 (15.8)	-22.2 (12.9)	0.031¹
NRS-Itch	8.0 (1.3)	4.8 (3.4)	-3.2 (2.9)	0.098 ²
BSA	89.4 (8.2)	37.9 (35.1)	-51.5 (36.1)	0.063 ¹
Head	8.8 (0.3)	4.2 (4.1)	-4.7 (4.1)	0.106 ²
Upper Limbs	16.8 (1.3)	7.7 (7.4)	-9.1 (8.1)	0.106 ²
Trunk	29.7 (4.9)	14.0 (13.0)	-15.6 (14.4)	0.106 ²
Lower Limbs	34.2 (2.5)	12.0 (13.6)	-22.2 (12.3)	0.059 ²
Skindex-16	83.2 (6.9)	47.0 (25.5)	-36.2 (29.4)	0.063 ¹
DLQI	21.3 (9.2)	8.7 (6.2)	-12.7 (10.0)	0.063 ¹

1. Wilcoxon signed rank exact test. 2. Wilcoxon signed rank test with continuity correction

*PGA baseline assessed at Week 4

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Y VENEREOLOGÍA



FUNDACIÓN
PIEL SANA
ACADEMIA ESPAÑOLA
DE DERMATOLOGÍA
Y VENEREOLOGÍA

Con el patrocinio de:



ucb