

Greater Achievement of Minimal Disease Activity With Abrocitinib Versus Dupilumab: An AHEAD Treat-to-Target Analysis of Patients With Moderate-to-Severe Atopic Dermatitis

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Introduction

Background

- Atopic dermatitis (AD) is a common inflammatory skin disease characterized by recurrent eczematous skin lesions and pruritus with heterogenous skin manifestations, symptoms, and severity^{1,2}
- Abrocitinib is an oral JAK1 selective inhibitor approved for the treatment of adults and adolescents with moderate-to-severe AD³⁻⁵
- The AHEAD (Aiming High in Eczema/Atopic Dermatitis) treat-to-target (T2T) approach introduces the concept of minimal disease activity (MDA) as the overarching goal, achieved when optimal targets are met across selected clinician- and patient-reported outcomes⁶
- Abrocitinib is well suited for integration into T2T protocols, potentially enabling more patients to attain MDA

Objective

- To assess the efficacy of abrocitinib versus dupilumab using the AHEAD T2T criteria

JAK1, Janus kinase 1.

1. Langan SM et al. *Lancet*. 2020;396:345-360. 2. Afshari M et al. *Front Immunol*. 2024;15:1361005. 3. Cibinqo 100 mg film-coated tablets. Summary of product characteristics. Kent: Pfizer, Limited; 2025. 4. European Medicines Agency, Cibinqo (abrocitinib). Summary of product characteristics. Pfizer Europe MA EEIG; 2024. 5. Cibinqo (abrocitinib) tablets, for oral use. Prescribing information. Pfizer Inc.; 2023. 6. Silverberg JI et al. *J Eur Acad Dermatol Venereol*. 2024;38:2139-2148.

AHEAD Treat-to-Target Endpoints

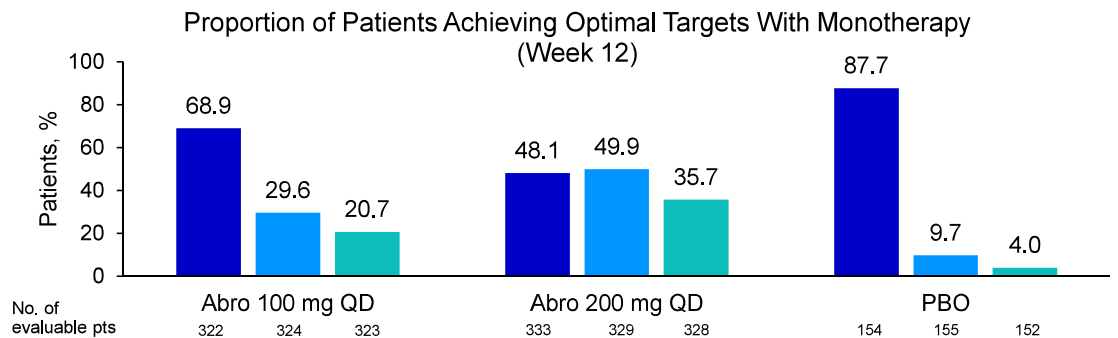
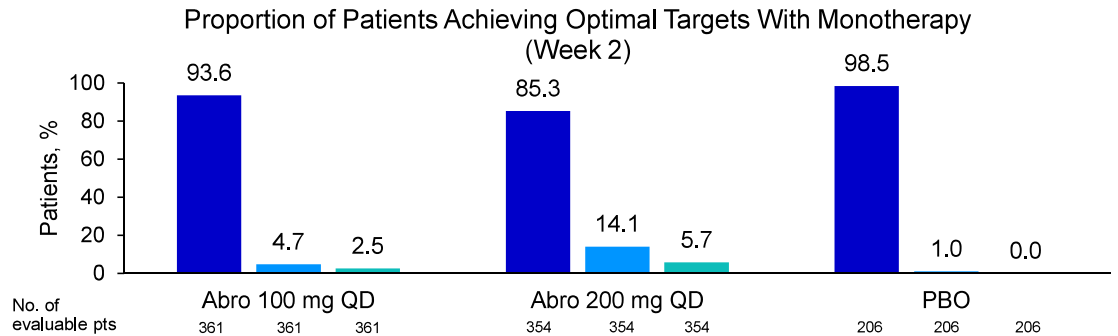
Outcome measure	Moderate target	Optimal target
Patient-reported outcomes		
PP-NRS	≥4-point improvement	PP-NRS <2
POEM	≥4-point improvement	POEM ≤2
POEM	≥3-point improvement in item 2 (sleep)	POEM (item 2) ≤2
HADS-A/D	HADS-A <11 OR HADS-D <11	HADS-A <8 AND HADS-D <8
PSAAD	≥3-point improvement in item 2 (pain)	PSAAD (item 2) <2
DLQI/cDLQI	≥4-point improvement (reduction)	DLQI/cDLQI <2
Clinical endpoints		
EASI	EASI-75 or EASI ≤7	EASI-90 OR EASI ≤3
SCORAD	SCORAD-50 or SCORAD ≤24	SCORAD-75 OR SCORAD ≤10
IGA, BSA	IGA ≤2 AND 50% improvement in BSA	IGA 0/1 AND BSA ≤2%

- This analysis included adult and adolescent patients ≥12 years with moderate-to-severe AD receiving abrocitinib monotherapy (100 or 200 mg) or placebo in a phase 2b (NCT02780167)¹ and the phase 3 JADE MONO-1/2 trials (NCT03349060/NCT03575871)^{2,3}
- Patients receiving abrocitinib (100 or 200 mg) or dupilumab 300 mg in combination with topical therapy in the JADE COMPARE/TEEN trials (NCT03720470/NCT03796676)^{4,5} were also included
- MDA was defined as the achievement of optimal targets for ≥1 clinical endpoint and ≥1 patient-reported outcome (PRO)
 - Treatment targets should be reviewed every 3-6 months and if targets are not achieved, treatment escalation or modification should be considered⁶

AD, atopic dermatitis; BSA, body surface area; cDLQI, Children's Dermatology Life Quality Index; DLQI, Dermatology Life Quality Index; EASI, Eczema Area and Severity Index; EASI-75, 75% improvement in EASI; EASI-90, 90% improvement in EASI; HADS-A, Hospital Anxiety and Depression Scale (Anxiety); HADS-D, Hospital Anxiety and Depression Scale (Depression); IGA, Investigator's Global Assessment; MDA, minimal disease activity; POEM, Patient-Oriented Eczema Measure; PP-NRS, Peak Pruritus Numerical Rating Score; PSAAD, Pruritus and Symptoms Assessment for Atopic Dermatitis; SCORAD, SCORing Atopic Dermatitis; SCORAD-50, 50% improvement in SCORAD; SCORAD-75, 75% improvement in SCORAD.
 1. Gooderham MJ et al. *JAMA Dermatol*, 2019;155:1371-1379. 2. Simpson EL et al. *Lancet*, 2020;396:255-266. 3. Silverberg JI et al. *JAMA Dermatol*, 2020;156:863-873. 4. Bieber T et al. *N Engl J Med*, 2021;384:1101-1112. 5. Eichenfield LF. *JAMA Dermatol*, 2021;157:1165-1173. 6. Silverberg JI et al. *J Eur Acad Dermatol Venereol*. 2024;38:2139-2148.

Achievement of MDA in Patients Receiving Monotherapy

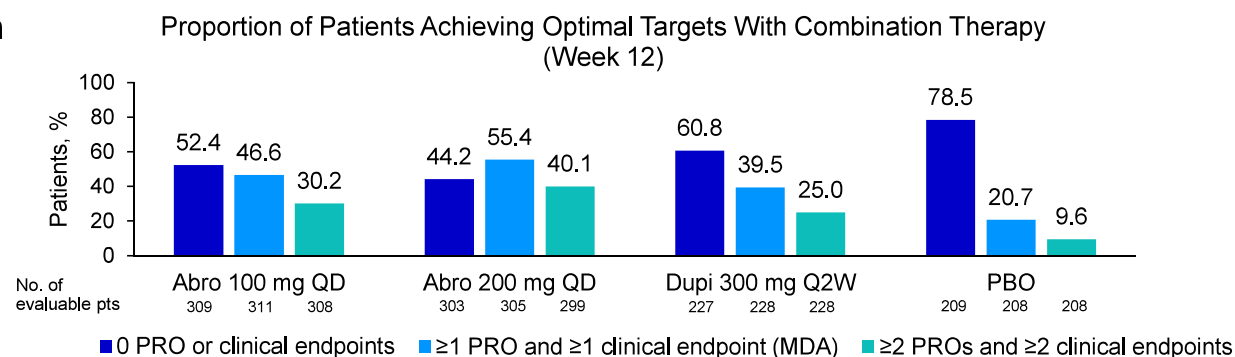
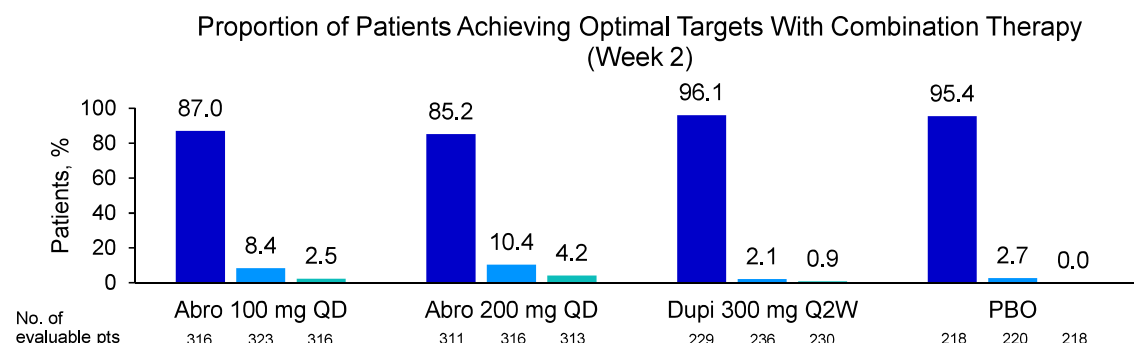
- Following 2 weeks of monotherapy treatment, 4.7% and 14.1% of patients receiving abrocitinib 100 or 200 mg, respectively, achieved MDA compared with 1.0% of patients receiving placebo
- By Week 12, rates of MDA increased to 29.6% and 49.9% in patients receiving abrocitinib 100 or 200 mg, respectively
- Notable proportions of patients achieved ≥ 2 PROs and ≥ 2 clinical endpoints in both the abrocitinib 100 mg (20.7%) and abrocitinib 200 mg (35.7%) groups
 - These results indicate that among patients receiving abrocitinib monotherapy who achieve MDA, most achieve optimal targets for multiple PROs and clinical endpoints



■ 0 PRO or clinical endpoints ■ ≥ 1 PRO and ≥ 1 clinical endpoint (MDA) ■ ≥ 2 PROs and ≥ 2 clinical endpoints

Achievement of MDA in Patients Receiving Combination Therapy

- Following 2 weeks of combination therapy, greater proportions of patients receiving abrocitinib 100 or 200 mg (8.4% and 10.4%) achieved MDA compared with those receiving dupilumab 300 mg (2.1%) or placebo (2.7%)
 - By Week 12, rates of MDA increased to 46.6% and 55.4% of patients receiving abrocitinib 100 or 200 mg monotherapy, respectively, and remained greater than in those receiving dupilumab (39.5%) or placebo (20.7%)
- Similar to patients receiving monotherapy, notable proportions of patients who achieved MDA achieved optimal targets for ≥ 2 PROs and clinical endpoints



- This analysis highlights the efficacy of abrocitinib in achieving MDA in patients with AD using a structured T2T approach following 12 weeks of consistent treatment with either monotherapy or concomitant topical therapy
- Rates of MDA were greater in patients receiving abrocitinib vs dupilumab, and most achieved optimal targets across multiple PROs and clinical endpoints simultaneously