



This is a repository copy of *Dupilumab reduces absenteeism in patients with moderate to severe atopic dermatitis: Pooled results from the LIBERTY AD SOLO clinical trials.*

White Rose Research Online URL for this paper:
<https://eprints.whiterose.ac.uk/166195/>

Version: Published Version

Article:

de Bruin-Weller, M., Simpson, E.L., Cork, M. orcid.org/0000-0003-4428-2428 et al. (6 more authors) (2020) Dupilumab reduces absenteeism in patients with moderate to severe atopic dermatitis: Pooled results from the LIBERTY AD SOLO clinical trials. *Journal of the American Academy of Dermatology*, 83 (5). pp. 1499-1501. ISSN 0190-9622

<https://doi.org/10.1016/j.jaad.2020.05.142>

Reuse

This article is distributed under the terms of the Creative Commons Attribution (CC BY) licence. This licence allows you to distribute, remix, tweak, and build upon the work, even commercially, as long as you credit the authors for the original work. More information and the full terms of the licence here:
<https://creativecommons.org/licenses/>

Takedown

If you consider content in White Rose Research Online to be in breach of UK law, please notify us by emailing eprints@whiterose.ac.uk including the URL of the record and the reason for the withdrawal request.



eprints@whiterose.ac.uk
<https://eprints.whiterose.ac.uk/>

LETTER

RESEARCH LETTER

Dupilumab reduces absenteeism in patients with moderate to severe atopic dermatitis: Pooled results from the LIBERTY AD SOLO clinical trials

To the Editor: Moderate to severe atopic dermatitis (AD) negatively affects daily functioning, quality of life (QoL), and work productivity,¹ more so for patients with inadequate symptom control.² The SOLO 1 and 2 phase 3 clinical trials showed that dupilumab versus placebo significantly improved signs, symptoms, and QoL in patients with moderate to severe AD, with an acceptable safety profile.³ This study evaluated the impact of dupilumab on work/school productivity using a pooled analysis of the SOLO trials.

The designs and primary findings from the SOLO trials have been reported previously.³ Adults with moderate to severe AD inadequately controlled by topical treatments received dupilumab (300 mg subcutaneously weekly or every 2 weeks) or placebo (subcutaneously weekly) over 16 weeks. We evaluated the mean number of missed days from work/school (absenteeism) and related costs in patients reporting full-time employment or school (≥ 4 days/week) and in patient subgroups categorized by baseline symptom severity, including Investigator's

Global Assessment of 3 or 4, Peak Pruritus Numerical Rating Scale score of less than 7 or 7 or greater, and Dermatology Life Quality Index of less than or equal to 10 or greater than 10. Duration-adjusted annualized absenteeism rate at week 16 was estimated using Poisson regression with treatment, region, and baseline Investigator's Global Assessment strata as fixed factors and the log value of assessment days up to weeks 4, 8, 12, or 16 as offset variables. Annual productivity costs for a hypothetical cohort of 10,000 treated patients were calculated using daily labor or employee compensation costs (2018 European Union⁴ and US⁵ rates).

Most (72%) SOLO participants reported full-time work/school commitments, with similar baseline characteristics among groups (Table I). Absenteeism rates first collected at week 4 showed a difference between dupilumab and placebo groups (mean [standard error] missed days per patient-year: placebo, 12.1 [0.76]; dupilumab every 2 weeks, 5.4 [0.49]; dupilumab every week, 7.0 [0.55]). Annualized absenteeism rates by week 16 were significantly lower with dupilumab (Table II), with 5.6 (every 2 weeks) and 4.4 (every week) fewer missed days per patient-year versus placebo. This finding persisted in analyzed subgroups, with greater reductions in missed days among those with greater baseline AD severity (Table II). Reduced

Table I. Baseline characteristics of SOLO participants with full-time work/school commitments (full analytic sample)

| Baseline characteristic | Placebo (n = 319) | Dupilumab 300 mg every 2 weeks (n = 332) | Dupilumab 300 mg every week (n = 346) |
|---------------------------------------------------------|-------------------|------------------------------------------|---------------------------------------|
| Age, y, median (Q1-Q3), range | 35 (26-46), 18-64 | 35 (26-44), 18-77 | 36 (26-47), 18-80 |
| Male sex, n (%) | 185 (58) | 206 (62) | 228 (66) |
| BMI, kg/m ² , median (Q1-Q3) | 25.1 (22.6-29.0) | 25.2 (22.7-28.7) | 25.3 (22.1-28.7) |
| Duration of AD, y, median (Q1-Q3) | 27 (19-38) | 26 (18-38) | 25 (17-37) |
| Total EASI, median (Q1-Q3) | 29 (22-42) | 29 (21-40) | 29 (22-43) |
| IGA score, n (%) | | | |
| 3 (moderate) | 173 (54.2) | 174 (52.4) | 180 (52.0) |
| 4 (severe) | 146 (45.8) | 158 (47.6) | 166 (48.0) |
| Weekly averaged Peak Pruritus NRS score, median (Q1-Q3) | 7.7 (6.4-8.7) | 7.7 (6.3-8.8) | 7.6 (6.3-8.6) |
| DLQI, median (Q1-Q3) | 14 (8-20) | 14 (9-20) | 15 (9-21) |

AD, Atopic dermatitis; BMI, body mass index; DLQI, Dermatology Life Quality Index; EASI, Eczema Area and Severity Index; IGA, Investigator's Global Assessment; NRS, Numerical Rating Scale; Q1, quartile 1; Q3, quartile 3.

Table II. Duration-adjusted annualized absenteeism rate* per patient-year by baseline disease severity at week 16

| | Placebo | Dupilumab 300 mg every 2 weeks | Dupilumab 300 mg every week |
|----------------------------------------------|-------------|-----------------------------------|--------------------------------|
| Full analytic sample | | | |
| n | 319 | 332 | 346 |
| Absenteeism rate, days per patient-year (SE) | 9.1 (0.34) | 3.5 (0.20) [†] | 4.7 (0.23) [†] |
| 95% CI | 8.4-9.8 | 3.1-3.9 | 4.3-5.2 |
| IGA score of 3 | | | |
| n | 173 | 174 | 180 |
| Absenteeism rate, days per patient-year (SE) | 6.4 (0.36) | 3.5 (0.26) [†] | 3.6 (0.27) [†] |
| 95% CI | 5.7-7.1 | 3.0-4.0 | 3.1-4.1 |
| IGA score of 4 | | | |
| n | 146 | 158 | 166 |
| Absenteeism rate, days per patient-year (SE) | 11.3 (0.55) | 3.2 (0.28) [†] | 5.7 (0.37) [†] |
| 95% CI | 10.2-12.4 | 2.6-3.7 | 5.0-6.4 |
| Peak pruritus NRS of <7 | | | |
| n | 100 | 112 | 114 |
| Absenteeism rate, days per patient-year (SE) | 5.0 (0.46) | 1.3 (0.20) [†] | 3.2 (0.32) [‡] |
| 95% CI | 4.1-6.0 | 0.9-1.7 | 2.5-3.8 |
| Peak pruritus NRS of ≥7 | | | |
| n | 219 | 219 | 230 |
| Absenteeism rate, days per patient-year (SE) | 11.0 (0.44) | 4.8 (0.29) [†] | 5.7 (0.31) [†] |
| 95% CI | 10.1-11.8 | 4.2-5.4 | 5.1-6.3 |
| DLQI of ≤10 | | | |
| n | 112 | 108 | 111 |
| Absenteeism rate, days per patient-year (SE) | 1.7 (0.26) | 0.6 (0.14) [†] | 0.9 (0.18) [‡] |
| 95% CI | 1.2-2.2 | 0.3-0.8 | 0.5-1.2 |
| DLQI of >10 | | | |
| n | 207 | 224 | 235 |
| Absenteeism rate, days per patient-year (SE) | 13.0 (0.49) | 5.0 (0.29) [†] | 6.5 (0.33) [†] |
| 95% CI | 12.0-13.9 | 4.4-5.5 | 5.9-7.2 |

Higher scores indicate worse signs (IGA), symptoms (Peak Pruritus NRS), and negative impact on quality of life (DLQI).

CI, Confidence interval; DLQI, Dermatology Life Quality Index; IGA, Investigator's Global Assessment; NRS, Numerical Rating Scale; SE, standard error.

*Estimated using Poisson regression with treatment, region, and baseline IGA strata as fixed factors and the log value of assessment days up to weeks 4, 8, 12, or 16 used as offset variables.

[†] $P < .0001$ versus placebo.

[‡] $P < .01$ versus placebo.

absenteeism resulted in lower productivity costs with dupilumab. Annual productivity costs avoided for the hypothetical cohort (based on annualized reduction of 5.6 days per patient-year with dupilumab every 2 weeks) were US\$16.2 million (United States), €15.3 million (Germany), €14.1 million (France), €12.6 million (Italy), £10.1 million (United Kingdom), and €9.5 million (Spain).

Dupilumab, thus, provided significant reductions in work/school absenteeism with associated cost savings versus placebo. Initial absenteeism and annualized reductions were greater with more severe AD and worse QoL at baseline.

Our findings may be different from a real-world population because of the controlled nature of clinical trials. The 16-week duration-adjusted analysis was extrapolated to annual estimates; the effect

of long-term treatment would require further investigation. There was no baseline assessment before treatment administration, with absenteeism rates first being collected at week 4; nonetheless, subsequent productivity outcomes were markedly different. Absenteeism rates were pooled across work/school settings and countries, with inherent variability that may affect generalizability.

The productivity burden of AD on patients with inadequately controlled disease is substantial but may be ameliorated with dupilumab treatment.

The authors would like to thank Qiuyue Chen for contributions to this work. Medical writing/editorial assistance was provided by Grace Richmond, PhD, of Excerpta Medica, funded by Sanofi Genzyme and Regeneron Pharmaceuticals, Inc.

Marjolein de Bruin-Weller, MD, PhD,^a Eric L. Simpson, MD, MCR,^b Michael Cork, BSc, MB, PhD, FRCP,^c Zhen Chen, PhD, MS, MA,^d Jerome Msibid, MSc,^e Christine Taniou, MSc,^f Laurent Eckert, PhD,^e Abhijit Gadkari, PhD,^d and Gaëlle Bégo-Le Bagousse, MSc^e

From the University Medical Center Utrecht, Utrecht, the Netherlands^a; Department of Dermatology, Oregon Health & Science University, Portland, Oregon^b; Sheffield Dermatology Research, The University of Sheffield, Sheffield Children's Hospital, Sheffield, United Kingdom^c; Regeneron Pharmaceuticals, Inc, Tarrytown, New York^d; Sanofi, Chilly-Mazarin, France^e; and Altran Technologies, Vélizy-Villacoublay, France.^f

Funding sources: Supported by Sanofi and Regeneron Pharmaceuticals, Inc.

Disclosure: Dr de Bruin-Weller is a consultant/advisor for AbbVie, Eli Lilly, Pfizer, Regeneron Pharmaceuticals, Inc, Sanofi Genzyme, and UCB and has received grant/research support from Regeneron Pharmaceuticals and Sanofi Genzyme. Dr Simpson reports grants from Eli Lilly, Kyowa Hakko Kirin, LEO Pharma, Merck, Pfizer, and Regeneron Pharmaceuticals Inc and personal fees from Dermira, Eli Lilly, Galderma, LEO Pharma, Menlo Therapeutics, Novartis, Pfizer, Regeneron Pharmaceuticals Inc, Sanofi Genzyme, and Valeant. Dr Cork is an investigator and/or consultant for Astellas, Boots, Dermavant, Galapagos, Galderma, Hyphens, Johnson & Johnson, Kymab, LEO Pharma, L'Oréal, Menlo Therapeutics, Novartis, Oxagen, Pfizer, Procter & Gamble, Reckitt Benckiser, Regeneron Pharmaceuticals Inc, and Sanofi Genzyme. Dr Chen is an employee and shareholder of Regeneron Pharmaceuticals, Inc. Drs

Msibid, Eckert, and Bégo-Le Bagousse are employees and may hold stock and/or stock options in Sanofi. Dr Taniou is an employee of Altran Technology. Dr Gadkari was a full-time employee of Regeneron Pharmaceuticals Inc when this work was conducted, received salary and bonus from Regeneron Pharmaceuticals Inc, and is currently a full-time employee of Boehringer Ingelheim.

IRB approval status: Approved (PAR1-14-442, PAR1-14-443).

Reprints not available from the authors.

Correspondence to: Marjolein de Bruin-Weller, MD, PhD, University Medical Center Utrecht, Heidelberglaan 100, 3584 CX Utrecht, the Netherlands

E-mail: m.s.debruin-weller@umcutrecht.nl

REFERENCES

1. Eckert L, Gupta S, Amand C, Gadkari A, Mahajan P, Gelfand JM. Impact of atopic dermatitis on health-related quality of life and productivity in adults in the United States: an analysis using the National Health and Wellness Survey. *J Am Acad Dermatol*. 2017;77:274-279.
2. Ariëns LFM, van Nimwegen KJM, Shams M, et al. Economic burden of adult patients with moderate to severe atopic dermatitis indicated for systemic treatment. *Acta Derm Venereol*. 2019;99:762-768.
3. Simpson EL, Bieber T, Guttman-Yassky E, et al. Two phase 3 trials of dupilumab versus placebo in atopic dermatitis. *N Engl J Med*. 2016;375:2335-2348.
4. Eurostat Statistics Explained. Estimated hourly labour costs in the European Union. 2017. Available at: https://ec.europa.eu/eurostat/statistics-explained/index.php/Hourly_labour_costs. Accessed October 22, 2019.
5. US Bureau of Labor Statistics. Employer costs for employee compensation (June 2018). Available at: <https://www.bls.gov/news.release/pdf/eccec.pdf>. Accessed October 24, 2019.

<https://doi.org/10.1016/j.jaad.2020.05.142>