

Sleep disturbance in atopic dermatitis: prescribing patterns in a UK population-based study

Children and adolescents with atopic dermatitis and comorbid sleep disturbance are more likely to be prescribed sedating antihistamines, non-sedating antihistamines and melatonin.

Carsten Flohr^{1*}, Mandy Wan², Shona Cameron¹, Maciej Czachorowski⁴, Andrew Wildman⁵, Charlotte Curtis⁵, Melissa Watkins⁶

Affiliations:

¹ Paediatric and Population-Based Dermatology Research, St John's Institute of Dermatology, King's College London and Guy's and St Thomas' NHS Foundation Trust, London, UK. | ² Evelina London Children's Hospital, Guy's & St Thomas' NHS Foundation Trust, London, UK. | ³ Institute of Pharmaceutical Science, King's College London, London, UK. | ⁴ Pfizer Ltd., Tadworth, UK | ⁵ Momentum Data, Pendragon House, 65 London Road, St. Albans, UK | ⁶ Pfizer Inc., New York, USA

***Author for Correspondence**

Email: carsten.flohr@kcl.ac.uk

Conflict of Interest Statement: Carsten Flohr has received support from Pfizer for investigator-led research, as well as speaker, advisory and consultancy roles. Mandy Wan has received an honorarium from Vitaflo International Ltd. Shona Cameron has no conflict of interest to declare. Andrew Wildman and Charlotte Curtis are employees of Momentum Data, UK. Maciej Czachorowski and Melissa Watkins are employees and shareholders of Pfizer Inc.



Background & Aims

- **Sleep disturbance** is common in AD^{1,2} and impacts on daily life,³ including wider family.⁴
- Guidelines only recommend **short-term use of sedating antihistamines**.⁵ Limited evidence for **melatonin**.⁶
- There is little **real-world data** on **prescribing practices**.



Understanding prescribing in paediatric patients with AD and sleep disturbance is important.

1. Sugiyama A, *et al.* Acta Derm Venereol 2023; 103: adv12345.
2. Fishbein AB, *et al.* J Am Acad Dermatol 2018; 78(2): 336-41.
3. Gerner T, *et al.* J Eur Acad Dermatol Venereol 2021; 35(4): 948-57.
4. Ramirez FD, *et al.* JAMA Dermatol 2019; 155(5): 556-63.

4. National Institute for Health and Clinical Excellence. Eczema - atopic. Available from: <https://cks.nice.org.uk/topics/eczema-atopic/>
5. Chang YS, *et al.* A Randomized Clinical Trial. JAMA Pediatr 2016; 170(1): 35-42.

Methods



- **Children and adolescents (aged 2-<18 years)** registered within Clinical Practice Research Datalink (CPRD) Aurum database (01/03/2003 - 01/03/2023).

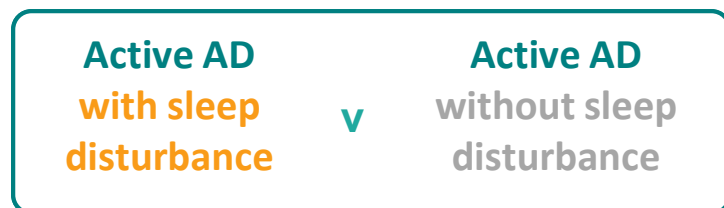


- **Active AD** cases were identified using at least one AD-specific clinical diagnosis code and 2 x AD treatment codes within 365 days.



- **Sleep disturbance (SD)** was a composite of sleep related clinical codes.

Overall



Subgroups:

- AD with comorbid asthma
- AD without comorbid asthma

Clinical and sociodemographic characteristics well balanced between groups



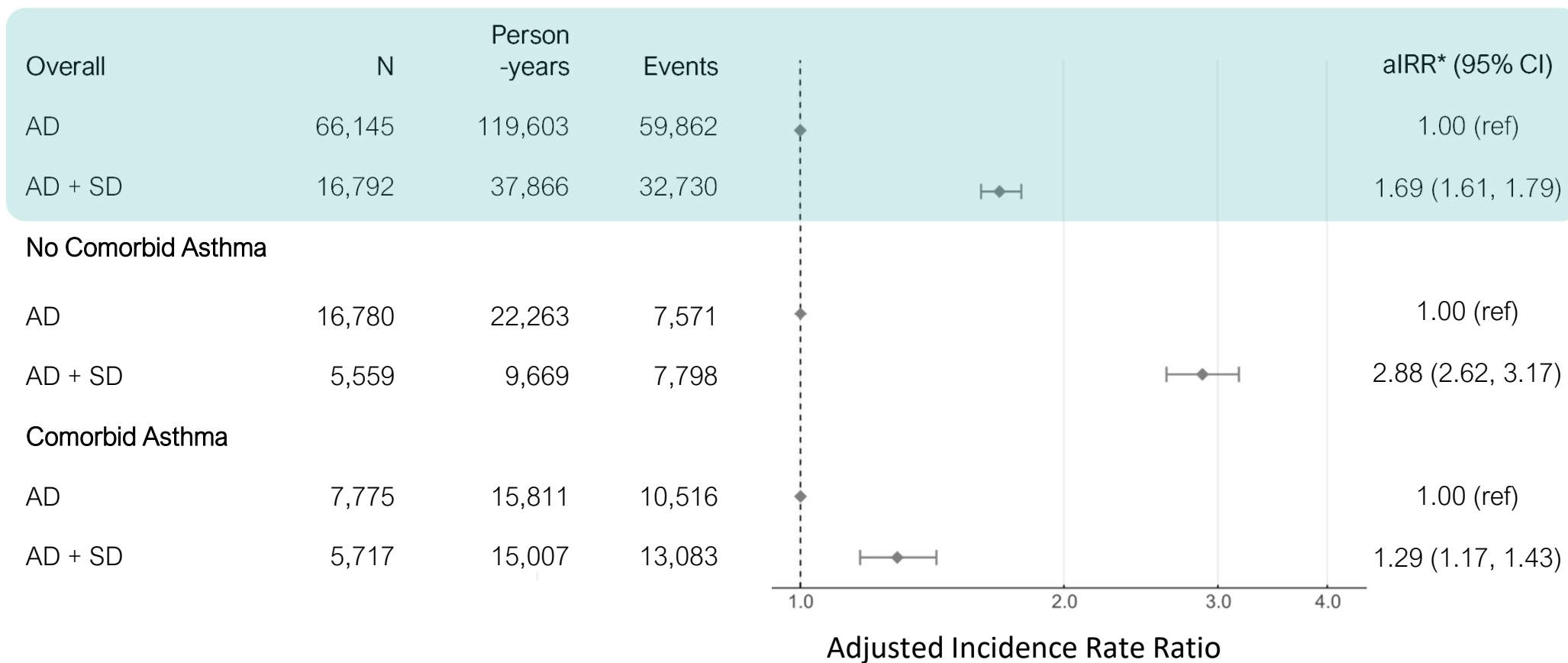
	Children with active AD	Matched controls	SMD ^a
N	643,012	2,546,802	
Age, median (IQR)	3.3 (2.0, 7.8)	3.2 (2.0, 7.6)	0.02
Follow-up, median (IQR)	1.34 (1.0, 2.0)	1.30 (0.9, 2.0)	0.06
Age groups			0.01
2-4	402,913 (63%)	1,606,937 (63%)	
5-11	157,668 (24%)	619,122 (24%)	
12-15	56,717 (9%)	221,309 (9%)	
16-17	25,714 (4%)	99,434 (4%)	
Sex			0.00
Male	325,635 (51%)	1,289,823 (51%)	
Female	317,377 (49%)	1,256,979 (49%)	
Ethnicity			0.00
White	332,557 (52%)	1,316,048 (52%)	
South Asian	69,189 (11%)	274,010 (11%)	
Black	41,565 (6.5%)	164,505 (6.5%)	
Mixed	19,933 (3.1%)	78,754 (3.1%)	
Other	9,416 (1.5%)	37,243 (1.5%)	
Chinese	3,687 (0.6%)	14,043 (0.6%)	
Unknown	166,665 (26%)	662,199 (26%)	

	Children with active AD	Matched controls	SMD ^a
Index of Multiple Deprivation (IMD)			0.00
1 (least)	122,544 (19%)	486,069 (19%)	
2	115,125 (18%)	456,395 (18%)	
3	115,165 (18%)	456,252 (18%)	
4	137,095 (21%)	542,900 (21%)	
5 (most)	152,662 (24%)	603,684 (24%)	
Unknown	421 (<0.1%)	1,502 (<0.1%)	
AD severity at baseline			
Mild	591,672	-	
Moderate	14,466	-	
Severe	36,874	-	
Allergic comorbidities	216,431 (8.5%)	125,298 (19%)	0.32
Allergic rhinitis	62,696 (2.5%)	40,690 (6.3%)	0.19
Asthma	103,714 (4.1%)	60,396 (9.4%)	0.21
Urticaria	57,989 (2.3%)	26,357 (4.1%)	0.10
Food allergy	13,224 (0.5%)	21,871 (3.4%)	0.21

IQR, Interquartile range;

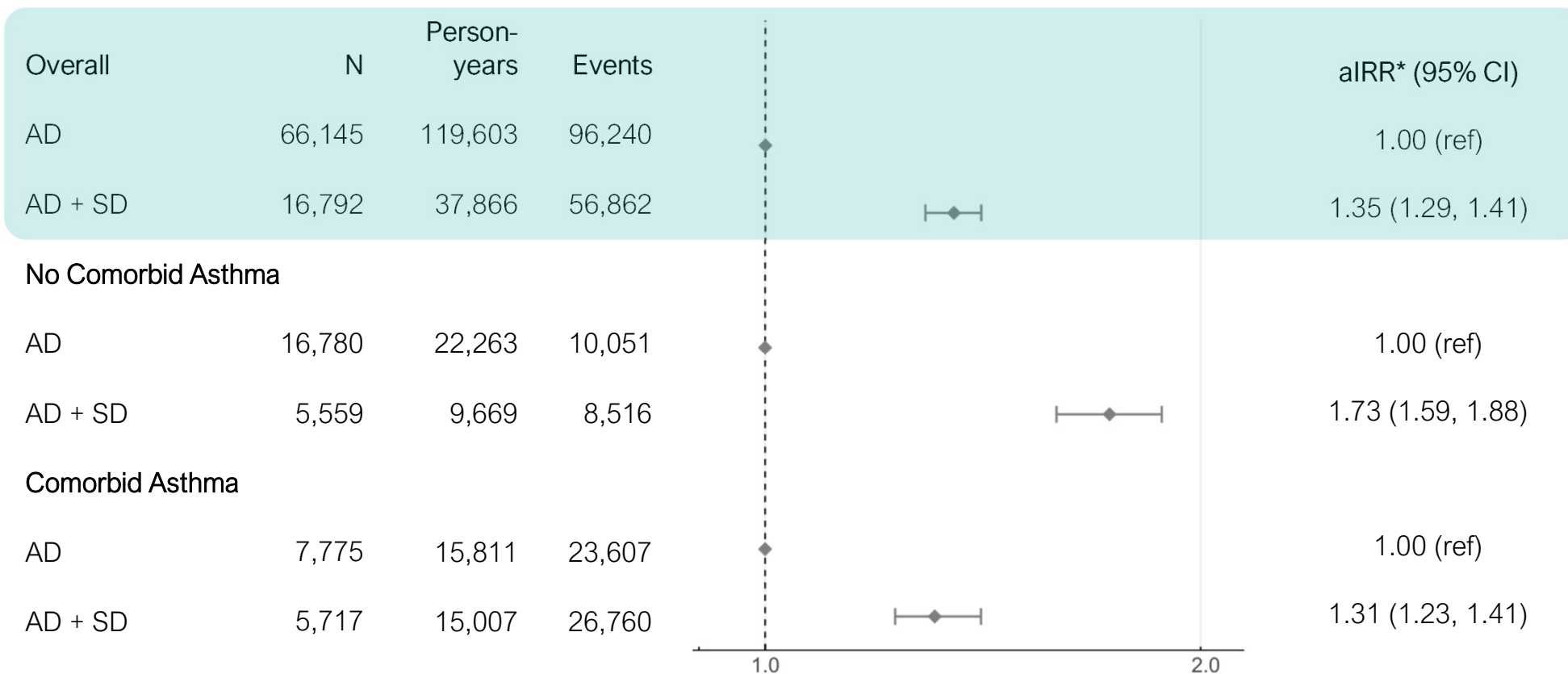
^aSMD, Standard Mean Difference, an estimate <0.1 indicates no meaningful difference between cases and controls.

Results: sedating antihistamines



*Adjusted for age, sex, ethnicity (white vs. non-white ethnicity) and allergic comorbidity (asthma, allergic rhinitis, urticaria, food allergy).

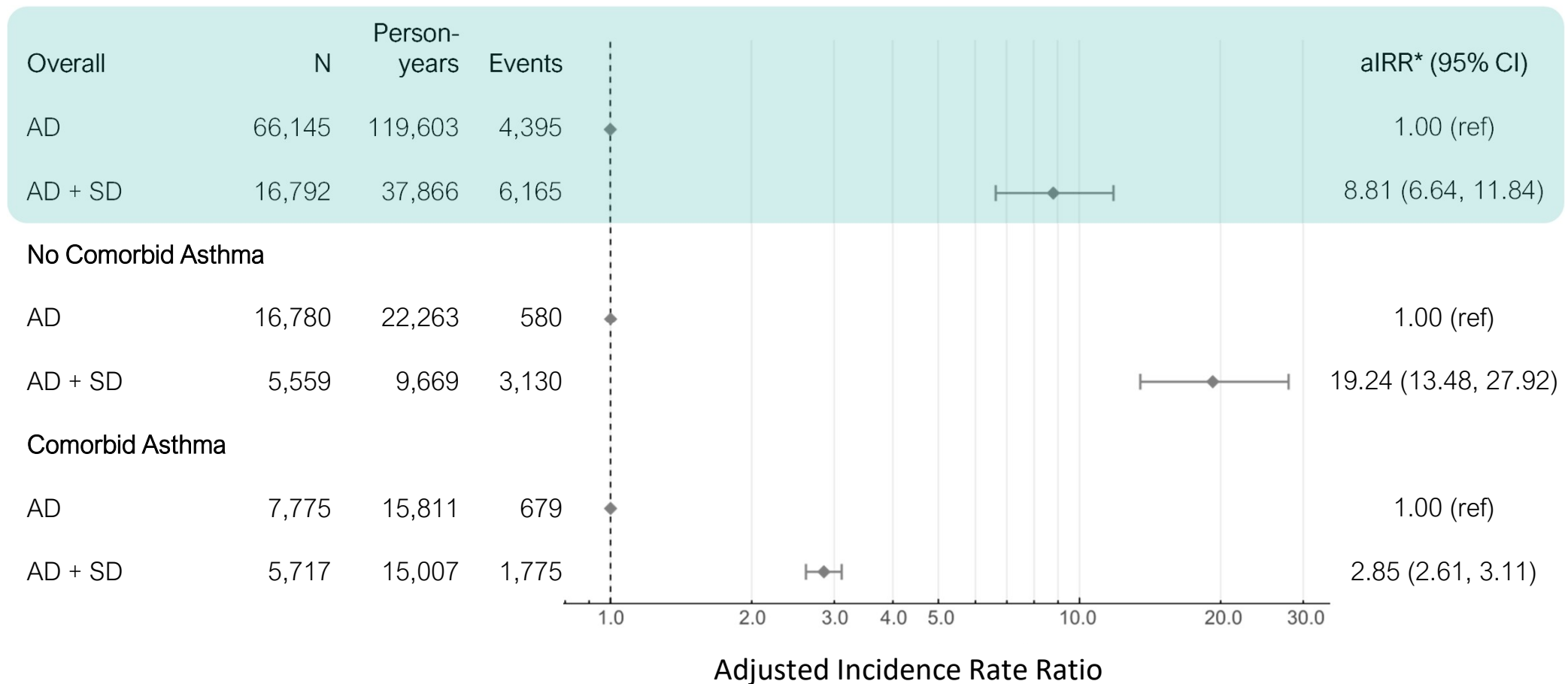
Results: non-sedating antihistamines



Adjusted Incidence Rate Ratio

*Adjusted for age, sex, ethnicity (white vs. non-white ethnicity) and allergic comorbidity (asthma, allergic rhinitis, urticaria, food allergy).

Results: melatonin

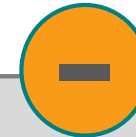


*Adjusted for age, sex, ethnicity (white vs. non-white ethnicity) and allergic comorbidity (asthma, allergic rhinitis, urticaria, food allergy).

Strengths & Limitations



- Population-based
- Validated AD case algorithm
- Detailed codes for sleep disturbances
- Large enough for subgroup analyses



- In UK, treatments also available without prescription.
- Unable to measure sleep duration.
- No clinical codes for AD-related sleep disturbance.

Conclusions



- Paediatric patients with **AD and sleep disturbance** are **more likely to be prescribed**:
 - Sedating antihistamines
 - Non-sedating antihistamines
 - Melatonin
- **Evidence** is needed on the most effective **treatment options for AD-related sleep disturbance** in paediatric patients.

Acknowledgements:

Project management support was provided by Emma Jones (emma.jones@momentumdata.co.uk) at Momentum Data, UK, and was funded by Pfizer Inc., New York, NY, USA, in accordance with Good Publication Practice (GPP 2022) guidelines (Ann Intern Med. 2022; 10.7362/M22-1460). Dr. Claire Feeney is also acknowledged for her contributions leading to study development.

This study is based in part on data from the Clinical Practice Research Datalink obtained under licence from the UK Medicines and Healthcare products Regulatory Agency. The data is provided by patients and collected by the NHS as part of their care and support. The interpretation and conclusions contained in this study are those of the author/s alone. Copyright © [2023], re-used with the permission of The Health & Social Care Information Centre. All rights reserved.

