Physical activity among children with asthma: cross-sectional analysis in the UK Millennium Cohort

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#### Abstract

Background Although beneficial for health and well-being, most children do not achieve recommended levels of physical activity. Evidence for children with asthma is mixed, with symptom severity rarely considered. This paper aimed to address this gap.

Methods We analysed cross-sectional associations between physical activity and parent-reported asthma symptoms and severity for 6497 UK Millennium Cohort Study 7-year-old-participants (3321 [49%] girls). Primary outcomes were daily moderate-to-vigorous physical activity (MVPA, minutes) and proportion of children achieving recommended minimum daily levels of 60 minutes of MVPA. Daily steps, sedentary time and total activity counts per minute (cpm) were recorded, as were parent-reported asthma symptoms, medications, and recent hospital admissions. Associations were investigated using quantile (continuous outcomes) and Poisson (binary outcomes) regression, adjusting for demographic, socioeconomic, health and environmental factors.

Results Neither asthma status nor severity were associated with MVPA; children recently hospitalised for asthma were less likely to achieve recommended daily MVPA (Risk Ratio, [95% CI]: 0.67 [0.44, 1.03]). Recent wheeze, current asthma and severe asthma symptoms were associated with fewer sedentary hours (difference in medians [95% CI]: -0.18 [-0.27, -0.08]; -0.14 [-0.24, -0.05]; -0.15 [-0.28, -0.02] respectively) and hospital admission with lower total activity (-48 cpm [-68, -28]).

Conclusion Children with asthma are as physically active as their asthma-free counterparts, while those recently hospitalised for asthma are less active. Qualitative studies are needed to understand the perceptions of children and families about physical activity following

hospital admission and to inform support and advice needed to maintain active lifestyles for children with asthma.

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#### Introduction

The physical and psychological benefits of physical activity are well known<sup>1,2</sup>. UK national guidelines recommend young people engage in at least 60 minutes moderate-to-vigorous physical activity (MVPA) daily<sup>3</sup>. Physical activity is likely to be particularly important for children with asthma since exercise training has been shown to reduce exercise-associated bronchconstriction<sup>4</sup> and might also counteract obesity and osteoporosis, which can be associated with inhaled and oral steroid treatments for asthma<sup>5,6</sup>.

Many UK children fail to meet recommended physical activity levels. Physical activity levels vary according to social and demographic factors, most notably gender and ethnicity<sup>7</sup>. Other factors, including health status, might also influence physical activity. Due to real or perceived limitation, children with chronic conditions might be at increased risk of poor engagement with physical activity; particularly those with conditions, such as asthma, where exercise can exacerbate symptoms<sup>8</sup>. It remains unclear, however, whether children with asthma are less likely to meet physical activity guidelines than children without asthma, and whether severity of asthma symptoms influences this risk.

A number of studies have considered the relationship between asthma and physical activity. Some have found children with asthma to be less physically active than those without<sup>9-11</sup>. Others, including a recent longitudinal study examining bidirectional associations<sup>12</sup>, have reported little difference<sup>13,14</sup> Generally, large population studies including children with asthma of varying severity have found similar levels of physical activity in those with and without asthma<sup>12,15,16</sup>. Studies reporting lower levels of physical activity in children with asthma compared to their peers without asthma have been small in size<sup>9-11</sup>, relied upon

enough to require follow-up in a hospital clinic<sup>9,10</sup>. To our knowledge, no studies have examined objective measures of physical activity or sedentary time in relation to asthma in a large nationally representative sample. Moreover, only a few studies have considered asthma severity<sup>11,14</sup> or symptom control<sup>17-19</sup> in relation to physical activity.

This study examines associations between parent-reported asthma and objectively-measured physical activity and sedentary time in a large, contemporary UK population-based sample. The association between asthma severity and physical activity is explored using indicators of symptom control, medication use and history of recent hospitalisation. We hypothesised that children with mild asthma symptoms would be no less active than their peers whereas for those with more severe asthma, physical activity levels might be lower.

#### Methods

## **Participants**

The Millennium Cohort Study (MCS) recruited 18, 818 children born in the UK between September 2000 and January 2002. Families with eligible children were identified through the Department of Work and Pensions Child Benefit system and selected on the basis of place of residence shortly after the participant child's birth. The sample is clustered at electoral ward level such that families living in the smaller UK countries, in disadvantaged areas and, in England, in areas with a high proportion of ethnic minorities, are over-represented<sup>20</sup>. Main carers for each child were first interviewed when the children were 9 months old and follow-up data were collected at successive surveys carried out throughout childhood. For 98% of children the main-carer respondent was the child's mother.

Of the original cohort, 14,043 (70%) participated in the fourth survey, at 7-8 years of age. The main care for each child participated in a computer-assisted home-based interview.

## Physical activity outcomes

At the fourth survey, children were invited to participate in the accelerometry study. Those whose parents consented were posted an accelerometer. Of the 13,219 children who agreed to participate in the accelerometer study, 7004, 6675, 6326, 5910, 5153, 4002 and 2244 had  $\geq 1$ ,  $\geq 2$ ,  $\geq 3$ ,  $\geq 4$ ,  $\geq 5$ ,  $\geq 6$  and  $\geq 7$  reliable days of data lasting at least 10 hours/day.<sup>21</sup>. This analysis was restricted to 6497 singleton children with at least two days with 10 or more hours of data each. Activity was measured using the Actigraph GT1M uni-axial accelerometer (Actigraph, Pensacola, Florida), initialised using ActiLife Lifestyle Monitoring

System software version 3.2.11 (Actigraph, Pensacola, Florida) and programmed to use a 15 second sampling epoch to record activity counts. Children were asked to wear their accelerometer on their right hip during all waking hours from the morning after they received it for seven consecutive days (except when in contact with water). Data were collected between May 2008 and August 2009 and downloaded using Actigraph software V.3.8.3 (Actigraph, Pensacola, Florida). Non-wear time, defined as consecutive zero activity counts  $\geq$  20 minutes, was excluded from analysis as were extreme count values  $\geq$ 11,715 counts per minute (cpm)<sup>22</sup>.

Threshold values for the accelerometer cpm were defined as <100 for sedentary time, and  $\leq 2240$ ,  $\leq 3840$ , and  $\geq 3841$  for light, moderate and vigorous physical activity, respectively<sup>23</sup>. Since the total valid recording time was not constant across days, total values for counts were standardised with reference to a standard day of equal duration for all children<sup>24</sup>. The primary outcome was mean daily minutes spent in MVPA. The proportion of children meeting the national recommendation of  $\geq 60$  minutes MVPA daily was also calculated. Secondary outcomes were mean daily hours spent sedentary, mean daily steps and total activity level (cpm).

Exposure variables: asthma and asthma severity

Exposure variables were based upon parental response to questions from the International Study of Asthma and Allergy in Childhood (ISAAC) core questionnaire for asthma age 6 – 7 years<sup>25</sup> and additional questions about medication use and hospital admission. All were measured at the 7-year survey.

#### **Asthma**

'Ever asthma' and 'recent wheeze' status was assigned according to the children's main-carers' responses to respectively 'has your child ever had asthma?', and 'has your child had wheezing or whistling in the chest in the last 12 months'. 'Current asthma' status was a derived variable requiring carers to affirm their child as ever having asthma and wheezing in the last 12 months or using asthma medications (based on reported use of 'any medications on a regular basis (including inhalers) that were prescribed by a doctor or hospital?' and including relievers or preventers identified via British National Formulary codes (Appendix S1)).

## Asthma severity

Children were categorised using the ISAAC definition of severe asthma and two further derived variables based upon reported asthma treatment or recent hospital admission for an asthma attack. To determine severity according to the ISAAC criteria carers were asked 'how many attacks of wheezing has your child had in the last 12 months?' (none, 1 to 3, 4 to 12, or > 12). They were then asked, 'in the last 12 months, how often on average has your child's sleep had been disturbed by wheezing?' (never, < 1 night/week, or ≥1 night/week), and 'in the last 12 months, has wheezing been bad enough to limit your child's speech to

one or two words at a time between breaths?' (yes or no). Those reporting  $\geq$  4attacks of wheeze per year, or  $\geq$  1nights/week of sleep disturbance due to wheeze, or wheeze affecting speech were classified as having severe asthma<sup>26</sup>. Those reporting prescription of any inhaled corticosteroid were compared with those who did not. Finally, carers reported whether the child had been admitted to hospital since last interview at age five years and reported the reason for the most serious admission. This information was used to categorise children as having experienced a recent hospital admission due to 'asthma or wheezing' (compared to those not admitted to hospital or for whom their most serious admission was for a condition other than asthma).

## Confounding factors

A number of potential common causes of asthma and physical activity were adjusted for a priori, as outlined in the directed acyclic graph (DAG)<sup>27</sup> (shown in Figure S1). Most were based on information collected from the main carer when the child was aged 7 years: child's sex, ethnicity, socioeconomic status (main carer's last employment or partner's if higher), number of siblings living in the household, country of residence at interview (England, Scotland, Northern Ireland or Wales) and household tobacco smoke exposure. Body mass index (BMI) was calculated for each child at age seven from measured weights and heights, which were converted to age- and sex-adjusted z scores using the UK 1990 growth reference<sup>28</sup>.

## Statistical analysis

The association between asthma status and physical activity was investigated before and after adjustment for confounding in multivariable models. As physical activity outcomes were positively skewed quantile regression was used to compare median differences between children categorised according to asthma status and asthma severity for the following outcomes, all measured daily: minutes of MVPA, total activity (cpm), hours of sedentary time, and total steps. Risk ratios (and 95% confidence intervals) were estimated in multivariable Poisson regression, in order to compare the proportion of children with and without asthma who met the recommended physical activity level of 60 minutes of MVPA daily. Survey weights were used to adjust for non-response and study sampling design<sup>20,24</sup>. A sensitivity analysis was performed excluding BMI since this might have a bidirectional association with physical activity level.

### Results

Of the 6497 children with valid accelerometer data, 6479 (99.6%) also had data from one or more of the asthma questions; of these, 979 (16.0%) reported ever receiving an asthma diagnosis, 743 (12.1%) reported wheeze within the last 12 months, and 512 (8.4%) reported current asthma. Severe asthma symptoms according to ISAAC criteria were reported by 323 (5.3%) children, 155 (2.5%) children reported being prescribed an inhaled corticosteroid, and 51 (0.7%) reported a hospital admission for asthma since last interviewed at 5 years. Median (IQR) values for activity outcomes were 594 cpm (504 to 697) daily total activity, 6.4 (5.8 to 7.1) hours spent sedentary, 60.5 (46.9 to 76.1) minutes spent in MVPA, and 10131 (8653 to 11794) steps daily. Including children with and without asthma the proportion

meeting recommended activity levels of 60 minutes MVPA daily was 50.9 (95% confidence interval (CI) 49.1 to 52.6%). Neither asthma prevalence nor sociodemographic nor other explanatory variables differed appreciably between the complete dataset of singleton children interviewed at 7 years and the smaller subset of children with activity data (Table 1).

Physical activity levels according to asthma status

Neither having ever received a diagnosis of asthma, nor current asthma were associated with MVPA (Table 2). Children reported to have experienced wheeze within the last 12 months recorded more daily minutes of MVPA (difference in medians, [95% CI]: 3.8, [1.0, to 6.5]) but this association was attenuated and did not remain significant after adjustment (0.5, [-1.6 to 2.6]) (Table 2). The proportion of children attaining daily recommended physical activity levels was low regardless of asthma status (Table 3). Of children reported to have ever received a diagnosis of asthma, 53.4% (95% CI: 49.4% to 57.4%) met recommendations compared to 50.3 (48.6 to 52.0) of children reported never to have been diagnosed with asthma. Similar patterns were observed for reported recent wheeze and current asthma.

Children reported to have experienced recent wheeze recorded greater total activity (cpm) and total daily steps, than those without recent reported wheeze (difference in medians, [95% CI]: 20, [2 to 38] and 329, [39 to 618] respectively), but these associations were attenuated and neither remained significant after adjustment (Table S1). Inclusion of socioeconomic status and gender in the adjusted model accounted for most of this attenuation. Children reported to have had recent wheeze also spent fewer daily hours

sedentary (difference in medians, [ 95% CI]: -0.21, [-0.30 to -0.11]) with little change after adjustment (-0.18, [-0.27 to -0.08]) (Table S1). Reports of ever asthma and current asthma were also associated with less daily sedentary time; however, both associations attenuated after adjustment with small associations with current asthma remaining (difference in medians, [95% CI]: -0.14, [-0.24 to -0.05]).

Physical activity levels according to asthma severity

Neither severe asthma symptoms (according to ISAAC criteria) nor prescription of inhaled corticosteroids was associated with daily MVPA. Children with a previous hospital admission for asthma reported fewer daily minutes of MVPA (difference in medians, [95% CI]: -7.1, [-13.6 to -0.5]) but this association was partly attenuated and no longer significant after adjustment (-5.1, [-13.9 to 3.7]) (Table 2). Proportions meeting national guidelines did not vary by severe asthma symptoms or prescribed inhaled corticosteroids (Table 3). Of those reporting an asthma admission since the previous interview the proportion meeting national guidelines was 36.0% (95% CI 19.8 to 52.2%) compared with 50.9% (49.4% to 52.5%) among those who did not report an admission (RR [95% CI]: 0.67 [0.44, 1.03]) (Table 3).

Children with a carer-reported hospital admission due to asthma since last interviewed were less active than those without such an admission (difference in medians, [95% CI]: -48 cpm, [-68 to -28]). (Table S1). In contrast, both children prescribed inhaled corticosteroids and those reported to have severe asthma symptoms spent fewer hours sedentary than their comparison groups. However both associations were attenuated after adjustment and only a small association with severe asthma symptoms remained (-0.15, [-0.28 to -0.02]).

Direction and size of n	nain effects were unaffe	ected by excluding BMI	I from the multivariable
model.			

#### Discussion

This study is the first to compare physical activity levels among children with and without asthma in a large prospective, representative UK-wide cohort, using objectively measured activity data from 6497 7 – 8-year-old children. Importantly, in contrast to cohorts drawn from clinical samples, the MCS sampling and study design did not select children with severe or poorly controlled asthma. Approximately half of all children who had ever received an asthma diagnosis did not meet recommended levels of 60 minutes daily MVPA; this is similar to the proportion for all children reported previously<sup>7</sup>. This proportion did not vary according to any of the asthma measures or by asthma severity, with the exception of children who hospitalised in the preceding two years for asthma. These children were less likely to meet UK guidelines and engaged in less total activity than those not reporting an admission. Sedentary time was less in children reporting recent wheeze, current asthma and severe asthma symptoms.

Our main finding that physical activity differs little between children with and without asthma differs from some previous studies which have reported reduced physical activity in children with asthma. These studies have for the most part recruited from paediatric outpatient clinics rather than from nationally representative population samples. Most used questionnaire-derived activity data and included small numbers of children, potentially with relatively severe or uncontrolled symptoms<sup>9,10</sup>. Of the few large population studies to have considered physical activity in children with asthma, many have relied upon questionnaire-derived activity data and unvalidated parental or self-reports of asthma status. These studies also found little evidence of reduced physical activity in association with

associations between asthma and sedentary time, in particular with computer use or television viewing<sup>15,16,29</sup>. Our findings are consistent, however, with a recent meta-analysis of studies objectively measuring physical activity and a report from the Avon Longitudinal Study of Parents and Children (ALSPAC); both found no evidence that physical activity levels differed between children with and without asthma<sup>30,31</sup>.

In our study the only exposure associated with lower physical activity was parent-reported hospital admission for asthma. Children previously admitted to hospital for asthma engaged in less total activity than those without a history of such an admission and were also less likely to meet national physical activity guidelines. Reduced physical activity was not associated with severity based on epidemiological criteria used in ISAAC or upon regular use of prescribed inhaled corticosteroids. Since a previous asthma attack is the strongest predictor of future attacks<sup>32</sup>, children who had experienced a hospital admission might be those most at risk and potentially fearful of future attacks. Moreover, where attacks occur due to poor asthma control, interval symptoms might restrict physical activity. This group included fewer children than the other indicators of severity. Whilst this might serve to identify children with the most severe asthma, the precision of outcome estimates for this group was low.

Generally, asthma severity is quantified according to the level of treatment needed to achieve symptom control. However associations between disease severity, medication prescription and clinical symptoms are complex since loss of symptom control can occur, because of under-prescribing, poor inhaler technique, non-adherence to medication, or severe or treatment-resistant disease. Studies which have compared physical activity in

controlled disease have reported conflicting results; some have found reduced physical activity in association with more severe disease<sup>11</sup>, or poorer control<sup>18</sup>, whilst others have found no association<sup>14,17</sup>. It is possible that previous hospital admission is a stronger influence upon physical activity than measures of medication or symptoms because it is a strong predictor of future attacks and this is something children with asthma (and their parents) wish to avoid.

While children reporting recent wheeze, current asthma and severe asthma symptoms were no more or less active than their peers, they did engage in less sedentary time. However these differences were small compared to overall sedentary time and are of uncertain clinical significance. It is possible that given the promotion of physical activity within asthma management guidelines<sup>33</sup> the associations seen between asthma status and less time spent sedentary might arise because parents of children with asthma might encourage light activity over sedentary time.

## Strengths and limitations

The findings from this study do not provide any evidence that children with asthma are less phsyically active than those without asthma, with the exception of the small proportion previously admitted to hospitalfor asthma who tended to be less active overall. Some studies have suggested a causal role for sedentary behaviour in asthma pathogenesis<sup>29,34</sup>, while others have suggested that the likelihood of receiving an asthma or wheeze diagnosis might be greater for physically active children<sup>15</sup>. Due to the cross-sectional nature of the MCS data we have analysed it is not possible to determine the direction of this relationship

and this was not the purpose of our study In this study we measured physical activity using a type of accelerometer that has been validated against heart rate monitoring, calorimetry and energy expenditure measured using doubly labelled water<sup>35</sup>. Specific activity count thresholds were calibrated for the device used and the participant age group<sup>23</sup>. Potential limitations of the study were underestimation of activity due to the need to remove the accelerometers during water-based or contact sports and risk of over interpretation because asthma symptoms were not recorded concurrently with physical activity.

Moreover, relying upon self-reported asthma status introduces the possibility for misclassification. Prevalence estimates reflect the methods used to estimate asthma prevalence as discussed by Sheikh et al<sup>36</sup> and as highlighted in our previous study comparing prevalence of parent reported with doctor-diagnosed asthma. This showed moderate to substantial agreement between parentally-reported and doctor-diagnosed asthma at this age in this cohort<sup>37</sup>. Nevertheless, the results of this study should be interpreted as a reflection of activity levels among children with parent-reported asthma.

### Clinical implications

Whilst, a comparable proportion of children with and without asthma failed to achieve the recommended physical activity levels, those who had experienced recent hospitalisation due to asthma were less likely to be moderately or vigorously active for 60 minutes or more each day. As shown previously<sup>7</sup>, half of UK children fail to meet recommended physical activity levels but the reasons for poor engagement might differ according to health status. Addressing fears surrounding exercise-induced asthma might be necessary to engage children with asthma with interventions designed to increase physical activity<sup>38</sup>. Physical training has been demonstrated not to worsen airway inflammation in children with

asthma<sup>39</sup>, exercise-induced bronchoconstriction is seen more often in those with poorly than well controlled asthma<sup>40</sup>, and greater physical activity has been shown to follow improved asthma control<sup>19</sup>. Therefore, providing children with asthma and their parents, teachers and healthcare professionals, with information about the benefits of gaining symptom control, and supporting them in achieving this would appear to be important components of any intervention specifically targeting children with asthma. Positively, children with asthma and severe asthma were less sedentary, possibly indicating that families may be taking steps to replace sedentary behaviours with light physical activity. They thus may be receptive to further increases in physical activity, where health professionals have identified greater physical activity as being appropriate and of benefit to that individual.

In conclusion we found no evidence to suggest that children in the general population with asthma - as reported by their parent - are less physically active than those without asthma and that they are slightly less sedentary. There is some suggestion from our findings that the very small percentage of children with asthma severe enough to require hospital admission are less active and thus less likely to meet national guidelines to achieve a minimum of an hour of moderate and vigorous active daily. When designing interventions to increase physical activity amongst children with asthma, children with an admission to hospital for an asthma attack might be at greatest risk of poor engagement. Further work is required to identify the real or perceived barriers to engagement in this and other at-risk groups. Lower sedentary behaviour among children with asthma indicate that families may be receptive to increasing physical activity, with the right information, guidance and support from health professionals.

#### References

- 1. Loprinzi PD, Cardinal BJ, Loprinzi KL, Lee H. Benefits and environmental determinants of physical activity in children and adolescents. Obes Facts 2012;5:597-610.
- 2. Lubans D, Richards J, Hillman C, Faulkner G, Beauchamp M, Nilsson M, Kelly P, Smith J, Raine L, Biddle S. Physical Activity for Cognitive and Mental Health in Youth: A Systematic Review of Mechanisms. Pediatrics 2016;38(3):e20161642.
- 3. Department of Health PA, Health Improvement and Protection. Start Active, Stay Active: A report on physical activity from the four home countries' Chief Medical Officers. 2011 https://www.gov.uk/government/uploads/system/uploads/attachment\_data/file/216370/d h\_128210.pdf. [Last accessed 17/12/2018].
- 4. Wanrooij VH, Willeboordse M, Dompeling E, van de Kant KD. Exercise training in children with asthma: a systematic review. Br J Sports Med 2014;48:1024-1031.
- 5. Hind K, Burrows M. Weight-bearing exercise and bone mineral accrual in children and adolescents: a review of controlled trials. Bone 2007;40:14-27.
- 6. Nogueira RC, Weeks BK, Beck BR. Exercise to improve pediatric bone and fat: a systematic review and meta-analysis. Med Sci Sports Exerc 2014;46:610-621.
- 7. Griffiths LJ, Cortina-Borja M, Sera F, Pouliou T, Geraci M, Rich C, Cole TJ, Law C, Joshi H, Ness AR, et al. How active are our children? Findings from the Millennium Cohort Study. BMJ Open 2013;3:e002893.
- 8. Bonini M and Palange P. Exercise-induced bronchoconstriction: new evidence in pathogenesis, diagnosis and treatment. Asthma Research and Practice 2015;1(2):s40733-015-0004-4.

- 9. Glazebrook C, McPherson AC, Macdonald IA, Swift JA, Ramsay C, Newbould R, Smyth A. Asthma as a barrier to children's physical activity: implications for body mass index and mental health. Pediatrics 2006;118:2443-2449.
- 10. Chiang LC, Huang JL, Fu LS. Physical activity and physical self-concept: comparison between children with and without asthma. J Adv Nurs 2006;54:653-662.
- 11. Lam KM, Yang YH, Wang LC, Chen SY, Gau BS, Chiang BL. Physical Activity in School-Aged Children with Asthma in an Urban City of Taiwan. Pediatr Neonatol 2016;57:333-337.
- 12. Cassim R, Dharmage SC, Koplin JJ, Milanzi E, Russell MA. Suspected asthma status and time spent in physical activity across multiple childhood age groups. Ann Allergy Asthma Immunol 2018;120:219-220.
- 13. van Gent R, van der Ent CK, van Essen-Zandvliet LE, Rovers MM, Kimpen JL, de Meer G, Klijn PH. No differences in physical activity in (un)diagnosed asthma and healthy controls. Pediatr Pulmonol 2007;42:1018-1023.
- 14. Sousa AW, Cabral AL, Martins MA, Carvalho CR. Daily physical activity in asthmatic children with distinct severities. J Asthma 2014;51:493-497.
- 15. Mitchell EA, Beasley R, Bjorksten B, Crane J, Garcia-Marcos L, Keil U, Group IPTS. The association between BMI, vigorous physical activity and television viewing and the risk of symptoms of asthma, rhinoconjunctivitis and eczema in children and adolescents: ISAAC Phase Three. Clin Exp Allergy 2013;43:73-84.
- 16. Corbo GM, Forastiere F, De Sario M, Brunetti L, Bonci E, Bugiani M, Chellini E, La Grutta S, Migliore E, Pistelli R, et al. Wheeze and asthma in children: associations with body mass index, sports, television viewing, and diet. Epidemiology 2008;19:747-755.

- 17. Matsunaga NY, Oliveira MS, Morcillo AM, Ribeiro JD, Ribeiro M, Toro A. Physical activity and asthma control level in children and adolescents. Respirology 2017;22:1643-1648.
- 18. Holderness H, Chin N, Ossip DJ, Fagnano M, Reznik M, Halterman JS. Physical activity, restrictions in activity, and body mass index among urban children with persistent asthma. Ann Allergy Asthma Immunol 2017;118:433-438.
- 19. Vahlkvist S, Inman MD, Pedersen S. Effect of asthma treatment on fitness, daily activity and body composition in children with asthma. Allergy 2010;65:1464-1471.
- 20. Plewis IC, L; Hawkes, D; hughes, G; Joshi, H. 2007. Millennium cohort study: technical report on sampling. 4th ed. London: Institute of Education, University of London.
- 21. Rich C, Geraci M, Griffiths L, Sera F, Dezateux C, Cortina-Borja M. Quality control methods in accelerometer data processing: defining minimum wear time. PLoS One 2013;8:e67206.
- 22. Geraci MR, C; Sera, F; Cortina-Borja, M: Griffiths, LJ; Dezateux, C. 2012. Technical report on accelerometry data processing in the Millennium Cohort Study. London: Institute of Child Health, University College London.
- 23. Pulsford RM, Cortina-Borja M, Rich C, Kinnafick FE, Dezateux C, Griffiths LJ. Actigraph accelerometer-defined boundaries for sedentary behaviour and physical activity intensities in 7 year old children. PLoS One 2011;6:e21822.
- 24. Griffiths LR, C; Geraci, M; Sera, F; Cortina\_Borja, M; Pouliou, T; Platt, L; Johnson, J; Dezateux, C. 2013. Technical report on the enhancement of Millennium Cohort Study data with accelerometer-derived measures of physical activity and sedentary behaviour in seven year olds, London: Institute of Child Health, University College London.

- 25. Asher MI, Keil U, Anderson HR, Beasley R, Crane J, Martinez F, Mitchell EA, Pearce N, Sibbald B, Stewart AW, et al. International Study of Asthma and Allergies in Childhood (ISAAC): rationale and methods. Eur Respir J 1995;8:483-491.
- 26. Lai CK, Beasley R, Crane J, Foliaki S, Shah J, Weiland S, International Study of A, Allergies in Childhood Phase Three Study G. Global variation in the prevalence and severity of asthma symptoms: phase three of the International Study of Asthma and Allergies in Childhood (ISAAC). Thorax 2009;64:476-483.
- 27. Textor J, Hardt J, Knuppel S. DAGitty: a graphical tool for analyzing causal diagrams. Epidemiology 2011;22:745.
- 28. Cole TJ, Freeman JV, Preece MA. British 1990 growth reference centiles for weight, height, body mass index and head circumference fitted by maximum penalized likelihood. Stat Med 1998;17:407-429.
- 29. Hark WT, Thompson WM, McLaughlin TE, Wheatley LM, Platts-Mills TA. Spontaneous sigh rates during sedentary activity: watching television vs reading. Ann Allergy Asthma Immunol 2005; 94:247-250.
- 30. Eijkemans M, Mommers M, Draaisma JM, Thijs C, Prins MH. Physical activity and asthma: a systematic review and meta-analysis. PLoS One 2012;7:e50775.
- 31. Jago R, Salway RE, Ness AR, Hamilton Shield JP, Ridd MJ, Henderson AJ.

  Associations between physical activity and asthma, eczema and obesity in children aged 12–16: an observational cohort study. BMJ Open 2019;9:e024858.
- 32. Miller MK, Lee JH, Miller DP, Wenzel SE, Group TS. Recent asthma exacerbations: a key predictor of future exacerbations. Respir Med 2007;101:481-489.

- 33. BTS/SIGN British guideline on the management of asthma: a national guideline 2016. https://www.brit-thoracic.org.uk/document-library/clinical-information/asthma/btssign-asthma-guideline-2016/ [Last accessed 17/12/2018]
- 34. Fredberg JJ, Inouye DS, Mijailovich SM, Butler JP. Perturbed equilibrium of myosin binding in airway smooth muscle and its implications in bronchospasm. Am J Respir Crit Care Med 1999; 159:959-967.
- 35. Puyau MR, Adolph AL, Vohra FA, Butte NF. Validation and calibration of physical activity monitors in children. Obes Res 2002;10:150-157.
- 36. Punekar YS and Sheikh A. Establishing the incidence and prevalence of clinician-diagnosed allergic conditions in children and adolescents using routinely collected data from general practices. Clin Exp Allergy 2009;39:1209–1216.
- 37. Griffiths LJ, Lyons RA, Bandyopadhyay A, Tingay KS, Walton S, Cortina-Borja M, Akbari A, Bedford H, Dezateux C. Childhood asthma prevalence: cross-sectional record linkage study comparing parent-reported wheeze with general practitioner-recorded asthma diagnoses from primary care electronic health records in Wales BMJ Open Respiratory Research 2018;5:e000260.
- 38. Yago R, Searle A, Henderson AJ, Turner KM. Designing a physical activity intervention for children with asthma: a qualitative study of the views of healthcare professionals, parents and children with asthma. BMJ Open 2017;7(3):e014020.
- 39. Moreira A, Delgado L, Haahtela T, Fonseca J, Moreira P, Lopes C, Mota J, Santos P, Rytila P, Castel-Branco MG. Physical training does not increase allergic inflammation in asthmatic children. Eur Respir J 2008;32:1570-1575.
- 40. Ostrom NK, Parsons JP, Eid NS, Craig TJ, Stoloff S, Hayden ML, Colice GL. Exercise-induced bronchospasm, asthma control, and obesity. Allergy Asthma Proc 2013;34:342-348.

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# Data availability:

All MCS data used in this analysis are available from UK Data Service, University of Essex and University of Manchester:

- http://doi.org/10.5255/UKDA-SN-8172-2
- http://doi.org/10.5255/UKDA-SN-6411-7
- http://doi.org/10.5255/UKDA-SN-7238-1

Table 1 Comparison of asthma status and sociodemographic characteristics between the whole cohort and the subset of singleton children with accelerometer data

Variable		All singleton children who participated in the age 7 survey  Percentages weighted according to survey weights			Subset singletons with physical activity data  Percentages weighted according to accelerometer  study weights		
		Male N (%)	Female N (%)	Total N	Male N (%)	Female N (%)	Total N
		6950 (51.4)	6731 (48.6)	13681	3176 (51.1)	3321 (48.9)	6497
Asthma ever	No	5572 (80.8)	5761 (86.6)	11333 (83.6)	2597 (81.4)	2903 (86.7)	5500 (84.0)
	Yes	1131 (19.2)	912 (13.4)	2243 (16.4)	570 (18.6)	409 (13.3)	979 (16.0)
Recent wheeze	No	5927 (86.0)	6019 (90.3)	11946 (88.1)	2736 (85.9)	3009 (90.0)	5745 (87.9)
Recent Wheele	Yes	989 (14.0)	667 (9.7)	1656 (11.9)	435 (14.1)	308 (10.0)	743 (12.1)
Current asthma	No	6181 (89.84)	6208 (93.1)	12389 (91.4)	2862 (90.0)	3103 (93.26)	5965 (91.6)
Current astrina	Yes	719 (10.2)	464 (6.9)	1183 (8.6)	304 (10.0)	208 (6.8)	512 (8.4)
Severe asthma	No	6446 (93.5)	6382 (95.7)	12828 (94.6)	2896 (93.9)	3179 (95.5)	6165 (94.7)
symptoms (ISAAC)	Yes	470 (6.5)	304 (4.3)	774 (5.4)	185 (6.1)	138 (4.5)	323 (5.3)
Prescribed inhaled	No	6742 (97.4)	6550 (98.0)	13292 (97.7)	3088 (97.4)	3241 (97.7)	6329 (97.5)
corticosteroids	Yes	174 (2.6)	129 (2.0)	303 (2.3)	2 (2.6)	73 (2.3)	155 (2.5)
Asthma admission	No	6832 (98.9)	6642 (99.4)	13474 (99.1)	3134 (99.1)	3301 (99.5)	6435 (99.3)
since age 5 years	Yes	84 (1.1)	44 (0.6)	128 (0.9)	35 (0.9)	16 (0.5)	51 (0.7)
Country	England	4451 (63.9)	4335 (64.6)	8786 (64.2)	2064 (64.7)	2140 (65.2)	4204 (64.9)
	Wales	1015 (14.8)	930 (13.6)	1945 (14.2)	450 (14.6)	448 (13.1)	898 (13.8)
	Scotland	809 (11.3)	790 (12.0)	1599 (11.6)	363 (11.1)	398 (11.9)	761 (11.5)
	Northern Ireland	675 (10.0)	676 (9.8)	1351 (9.9)	299 (9.6)	335 (9.8)	634 (9.7)
	Managerial and professional	3112 (46.2)	2978 (45.9)	6090 (46.1)	1743 (47.4)	1721 (45.0)	3464 (46.2)
Socio-economic	Intermediate	937 (13.5)	854 (13.4)	1791 (13.5)	418 (13.7)	430 (13.1)	848 (13.4)
Classification	Self-employed	650 (10.0	661 (10.1)	1311 (10.1)	268 (9.7)	308 (10.0)	576 (9.8)
(highest of main	Low supervisory and				205 (7.4)	237 (8.6)	442 (8.0)
carer and partner)	technical	509 (7.7)	497 (7.6)	1006 (7.7)			
	Semi-routine and	1487 (22.5)	1489 (22.9)	2976 (22.7)	480 (21.7)	560 (23.2)	1040 (22.5)
	routine						
Ethnicity	White	5830 (85.6)	5611 (85.3)	11441 (85.4)	2795 (85.0)	2916 (85.4)	5711 (85.2)
	Mixed	178 (3.1)	194 (3.4)	372 (3.2)	85 (3.5)	83 (3.0)	168 (3.2)
	Indian	179 (2.1)	162 (1.8)	341 (1.9)	61 (1.8)	78 (2.2)	139 (2.0)

	Pakistani/Bangladeshi	421 (4.5)	462 (5.0)	883 (4.7)	123 (5.4)	124 (4.9)	247 (5.1)
	Black	243 (3.5)	209 (3.1)	452 (3.3)	76 (3.2)	76 (2.9)	152 (3.0)
	Other	99 (1.3)	92 (1.5)	191 (1.4)	36 (1.2)	44 (1.7)	80 (1.4)
Number of	other children in	051 (10.5)	000 (10 4)	1/50/10 4)	224 (44.2)	255 (42.0)	/70 /11 /)
children in	household	851 (12.5)	808 (12.4)	1659 (12.4)	324 (11.3)	355 (12.0)	679 (11.6)
household	Only child	6099 (87.5)	5923 (87.6)	12022 (87.6)	2852 (88.7)	2966 (88.0)	5818 (88.4)
BMI (z-score)	NA (050/ 01)	0.40	0.32	0.36	0.39	0.30	0.35
	Mean (95% CI)	(0.37, 0.44)	(0.29, 0.36)	(0.34, 0.39)	(0.34, 0.44)	(0.25, 0.35)	(0.31, 0.38)
DNALka /no 2	Madian (IOD)	16.1	16.2	16.1	16.0	16.1	16.1
BMI kg/m²	Median (IQR)	(16.0, 16.2)	(16.1, 16.2)	(16.1, 16.2)	(15.9, 16.1)	(16.0, 16.2)	(16.0, 16.2)
Household	No	6068 (87.1)	5769 (85.7)	11837 (86.4)	2852 (87.4)	2935 (85.6)	5787 (86.5)
spoking exposure	Yes	847 (12.9)	912 (14.3)	1759 (13.6)	320 (12.6)	381 (14.4)	701 (13.5)

Missing data (main dataset, physical activity subset): asthma ever 105, 18; recent wheeze 79, 9; current asthma 109, 20; Severe asthma symptoms (ISAAC) 79, 9; hospital admission for asthma since age five years 79, 11; prescribed inhaled corticosteroids 86, 13; socioeconomic status 507, 127; Body mass index 221; 21 household smoking exposure 85, 9.

