## **SUPPLEMENTARY MATERIAL**

E-table 1. Definitions used in the questionnaires at SPAIS I–III.

Definitions	SPAIS I	SPAIS II	SPAIS III
Asthma	Ever having had	Ever having had	Ever having had
	parent- or self-	self-reported asthma	self-reported asthma
	reported asthma in	in combination with	in combination with
	combination with	having used inhaled	having used any
	having used inhaled	corticosteroid	inhaled asthma
	corticosteroid	treatment or having	treatment or having
	treatment or having	wheezing or	wheezing or
	wheezing or	whistling in the chest	whistling in the chest
	whistling in the chest	or having a	or having a
	or having a	respiratory infection	respiratory infection
	respiratory infection	that caused	that caused
	that caused	wheezing or	wheezing or
	wheezing or	whistling in the chest	whistling in the chest
	whistling in the chest	in the preceding	in the preceding
	in the preceding	year.	year.
	year.		
Wheeze	Having had	Having had	Having had
	wheezing or	wheezing or	wheezing or
	whistling in the chest	whistling in the chest	whistling in the chest
	at any time in the	at any time in the	at any time in the
	preceding year.	preceding year.	preceding year.
Rhinitis	Having had	Having had	Having had
	sneezing, nasal	sneezing, nasal	sneezing, nasal
	congestion, or	congestion, or	congestion, or
	rhinorrhoea during	rhinorrhoea during	rhinorrhoea during
	the preceding year,	the preceding year,	the preceding year,
	without having a	without having a	without having a
	cold.	cold.	cold.
Allergic symptoms	The subject's	Self-reported	Self-reported
to cat, dog or pollen	hypersensitivity to	hypersensitivity to	experience of ever
	cat, dog, or pollen,	cat, dog, or pollen,	having had allergic
	noticed and reported		symptoms to cat,
	by the parents.		dog,or pollen

E-table 2. Baseline characteristics of females and males in the study population and those lost to follow up, N = 959.

	Females			Males		
	SPAIS III	Lost to follow-up	p value	SPAIS III	Lost to follow-up	p value
	n = 273	n = 209		n = 218	n = 259	
FeNO <sub>0.1</sub> (ppb)	4.60	4.11		4.78	5.39	
	(4.19, 5.15)	(3.56, 4.73)	0.30	(4.13, 5.54)	(4.75, 6.12)	0.28
$FEV_1$						
(% predicted)	$94.69 \pm 0.58$	$95.33 \pm 0.84$	0.51	$95.06 \pm 0.76$	$95.04 \pm 0.65$	0.99
$FEV_1$						
(< -1.65 SD)						
(%)	7.0	11.1	0.11	8.3	7.4	0.73
BMI (kg/m <sup>2)</sup>	$19.7 \pm 0.18$	$20.3 \pm 0.23$	0.05	$19.8 \pm 0.20$	$19.9 \pm 0.19$	0.90
Height (cm)	$160.9 \pm 0.39$	$160.3 \pm 0.46$	0.35	$163.6 \pm 0.64$	$164.8 \pm 0.55$	0.18
Asthma (%)	8.1	10.1	0.45	8.7	8.1	0.81
Wheeze (%)	14.7	14.8	0.96	12.8	10.8	0.49
Rhinitis (%)	25.6	24.4	0.76	23.9	27.0	0.43
Family smoking						
(%)	30.4	42.6	0.006	24.8	33.6	0.04
Allergic						
symptoms to cat						
(%)	10.3	8.6	0.54	9.6	12.4	0.35
Allergic						
symptoms to						
dog (%)	6.2	4.3	0.36	6.4	4.3	0.29
Allergic						
symptoms to						
pollen (%)	16.9	15.8	0.76	20.6	17.0	0.31

BMI, body mass index; FeNO<sub>0.1</sub>, fractional exhaled nitric oxide measured at 100 mL/s; FEV<sub>1</sub>, forced expiratory volume in 1 second; ppb, parts per billion; SD, standard deviation. All results presented as percent (%) or mean  $\pm$  standard deviation or geometric mean and 95% confidence interval.

E-table 3. Development of reported symptoms between SPAIS I (age 12-15 years) and SPAIS II (age 16-19 years).

	Prevalence of symptoms at SPAIS I	Remission of symptoms between SPAIS I and II	Persistence of symptoms between SPAIS I and II	Incidence of symptoms between SPAIS I and II	Prevalence of symptoms at SPAIS II	†p value
Wheeze, (%)						
All subjects,	13.8	7.1	6.7	8.6	15.3	0.43
n = 491						
Females,	14.7	7.3	7.3	10.6	17.9	0.20
n = 273	10.0		6.0		11.0	0.71
Males, $n = 218$	12.8	6.9	6.0	6.0	11.9	0.71
Asthma, (%)	0.1	1.0	<i>-</i> 2	0.6	7.1	0.00
All subjects,	8.1	1.8	6.3	0.6	7.1	0.08
n = 491	0.1	1.1	7.0	0.4	7.2	0.22
Females, n= 273	8.1	1.1	7.0	0.4	7.3	0.32
Males, $n = 218$	8.3	2.8	5.5	0.9	6.4	0.16
Cat symptoms,	0.5	2.0	5.5	0.7	0.4	0.10
(%)						
All subjects,	10.2	0.4	9.8	5.4	15.2	< 0.001
n = 461						
Females,	10.5	0	10.5	6.2	16.7	< 0.001
n = 258						
Males, $n = 203$	9.9	1.0	8.9	4.4	13.3	0.03

 $<sup>\</sup>dagger p$  values compare prevalence rates at SPAIS I and II (McNemar's test). All results presented as percent (%).

E-table 4. Development of reported symptoms between SPAIS II (age 16–19 years) and SPAIS III (age 28–31 years).

	Prevalence of symptoms at SPAIS II	Remission of symptoms between SPAIS II and III	Persistence of symptoms between SPAIS II and III	Incidence of symptoms between SPAIS II and III	Prevalence of symptoms at SPAIS III	†p value
Wheeze,						
(%)						
All subjects, $n = 491$	15.3	7.5	7.7	10.6	18.3	0.11
Females, n = 273	17.9	8.1	9.9	11.7	21.6	0.17
Males, $n = 218$	11.9	6.9	5.0	9.2	14.2	0.40
Asthma,						
All subjects, n = 491	7.1	2.2	4.7	10.0	14.7	< 0.001
Females, n = 273	7.3	1.8	5.5	12.8	18.3	< 0.001
Males, $n = 218$	6.4	2.8	3.7	6.4	10.1	0.07
Cat						
symptoms, (%)						
All subjects, n = 461	15.2	2.2	13.0	9.5	22.6	< 0.001
Females, n = 258	16.7	3.5	13.2	8.9	22.1	0.01
Males, $n = 203$	13.3	0.5	12.8	10.3	23.2	< 0.001

 $<sup>^\</sup>dagger p$  values compare prevalence rates at SPAIS II and III (McNemar's test). All results presented as percent (%).

E-table 5. Sex differences in relation to baseline characteristics at SPAIS I (age 12–15 years) and data reported at SPAIS III (age 28–31 years).

	SPAIS I		SPAIS III				
	Males n=218	Females n=273	p value	Males n=218	Females n=273	p value	
FeNO <sub>0.1</sub> (ppb)	4.78 (4.13, 5.54)	4.60 (4.10, 5.15)	0.68	NR	NR		
FEV <sub>1</sub> (% predicted)	95.06± 11.19	$94.69 \pm 9.65$	0.70	NR	NR		
FEV <sub>1</sub> (< -1.65 SD), (%)	8.3	19 (7.0)	0.59	NR	NR		
Age (years)	$13.59 \pm 0.39$	$13.55 \pm 0.41$	0.29	NR	NR		
BMI $(kg/m^2)$	$19.83 \pm 2.96$	$19.74 \pm 2.97$	0.74	NR	NR		
Height (cm)	$163.6 \pm 9.49$	$160.9 \pm 6.43$	< 0.001	NR	NR		
Menarche, (%)	NR	72.5	NR	NR	NR		
Family asthma, (%)	25.7	33.7	0.06	NR	NR		
Family rhinitis, (%)	53.7	56.0	0.60	NR	NR		
Family smoking, (%)	24.8	30.4	0.17	NR	NR		
Current smoking, (%)	NR	NR		10.4	11.5	0.69	
Cat at home, (%)	28.4	25.6	0.49	13.4	17.7	0.19	
Dog at home, (%)	21.1	18.7	0.50	9.3	13.0	0.20	

BMI, body mass index; FeNO $_{0.1}$ , fractional exhaled nitric oxide measured at 100 mL/s; FEV $_1$ , forced expiratory volume in one second; NR, not relevant; ppb, parts per billion; SD, standard deviation. All results presented as percent (%) or mean  $\pm$  standard deviation or geometric mean and 95% confidence interval.

E-table 6. Sex differences in relation to prevalence of reported symptoms at SPAIS I (age 12–15 years), SPAIS II (age 16–19 years), and SPAIS III (age 28–31 years).

	SPAIS I			SPAIS II			SPAIS		
							III		
	Males	Females	p	Males	Females	p	Males	Females	p
	n = 218	n = 273	value	n = 218	n = 273	value	n = 218	n = 273	value
Wheeze, (%)	12.8	14.7	0.56	11.9	17.9	0.07	14.2	21.6	0.04
Rhinitis,	23.9	25.6	0.65	29.6	33.3	0.35	44.5	51.7	0.12
(%)									
	n = 203	n = 258		n = 203	n = 258		n = 203	n = 258	
Allergic symptoms to dog, (%)	5.9	6.6	0.77	8.4	8.5	0.95	11.8	14.7	0.36
Allergic symptoms to pollen, (%)	20.7	17.1	0.32	22.7	23.3	0.88	42.4	45.7	0.47

All results presented as percent (%).