

SUPPLEMENTARY MATERIAL

Table 1S. Sociodemographic and clinical characteristics

Characteristics	N (300)	
Age at reference date*		
Median (Q1, Q3)	50	39; 62
	N	%
Sex		
Male	83	27,7%
Female	217	72,3%
Working status		
Employed, full-time	151	50,3%
Employed, part-time	30	10,0%
Homemaker	29	9,7%
Student	13	4,3%
Unemployed	16	5,3%
Retired	45	15,0%
Disabled	7	2,3%
Other	9	3,0%
Comorbidities ¹		
Do not know/no comorbidities	25	8,3%
With comorbidities	275	91,7%
Presence of allergic sensitization (diagnosed by blood or skin testing)	150	50,0%
Allergic rhinitis	108	36,0%
Heart disease	17	5,7%
Anxiety	77	25,7%
Anaphylaxis	21	7,0%
Arthritis	28	9,3%
Aspirin sensitivity	25	8,3%
Atopic dermatitis/Eczema	46	15,3%
Chronic bronchitis	21	7,0%
Chronic Obstructive Pulmonary Disease (COPD)	9	3,0%
Chronic sinusitis	27	9,0%
Depression	51	17,0%
Diabetes	17	5,7%
Emphysema	5	1,7%
GERD (heartburn/reflux)	64	21,3%
Hypertension	52	17,3%
Chronic rhinosinusitis with Nasal Polyps	42	14,0%
Sleep Apnea	25	8,3%
Stroke	3	1,0%
Others	44	14,7%
Clinician-reported		

Age at initial diagnosis (years)		
Median (Q1, Q3)	30	14; 46
Years since initial diagnosis		
Median (Q1, Q3)	13	6; 29
Spirometry performed in the last year		
Yes	256	85,3%
No	44	14,7%
Asthma medications		
ICS (monotherapy)	12	4,0%
ICS + LABA	243	81,0%
LAMA	100	33,3%
LTRA	66	22,0%
SABA	10	3,3%
OCS	16	5%
Biologics	30	10%
Exacerbations experienced in the last year		
Number of patients with a severe exacerbation	116	38,7%
Number of patients with an exacerbation that resulted in hospitalization for asthma for ≥ 24 hours	21	7,0%
Number of severe exacerbations	N	%
Mean (SD)	0,8	1,31
Median (Q1, Q3)	0	0; 1
Min, max	0	6
Number of exacerbations requiring short term OCS treatment for at least three days AND/OR increase of OCS dosis ¹		
Mean (SD)	1,1	1,14
Median (Q1, Q3)	1	0; 1
Min, max	0	5
Number of exacerbations requiring visit to an emergency room (stay ≥ 24 hours), unplanned office visit or treatment with systemic corticosteroid ¹		
Mean (SD)	0,8	0,8
Median (Q1, Q3)	1	0; 1
Min, max	0	4
Number of exacerbations that resulted in hospitalization for asthma for ≥ 24 hours		
Mean (SD)	0,2	0,58
Median (Q1, Q3)	0	0; 0
Min, max	0	4

*Out of 300 patients, 2 (0.7%) were adolescents (<18 years old)

Abbreviations: GERD, gastroesophageal reflux disease; ICS, inhaled corticosteroid; LABA, long-acting beta-agonist; LAMA, long-acting muscarinic antagonist; LTRA, leukotriene receptor antagonist; OCS, oral corticosteroid; SABA, short-acting beta-agonist.

Table 2S. Asthma control definition

Asthma control levels	ACT	Exacerbations		ACQ-6	AIRQ
		Use of OCS or emergency visits /no planned in the last 12 months	Hospitalisations in the last 12 months		
Well controlled	≥20	0	0	≤0,7	≤1
Not well controlled	16-19	1	0	0,8-1,5	2-4
Very poorly controlled	≤15	≥2	≥1	>1,5	≥5

¹Among patients suffering any severe exacerbations

Table 3S. Performance characteristics of the AIRQ questionnaire for the Spanish cohort based on the participant's ACT score plus exacerbation history

AIRQ Models	Full Analysis Set (FAS)								
	LR+	LR-	Sensitivity	Specificity	Positive Predictive Value	Negative Predictive Value	AIC	AUC	Model Fit
Univariate Model 1¹									
AIRQ score cut-off									
≥ 1	2,37	0,04	97.7%	58.8%	85.7%	90.9%	238,467	78.2%	
≥ 2	6,33	0,12	89.3%	85.9%	94.1%	76.0%	200,992	87.6%	
≥ 3	16,01	0,26	75.3%	95.3%	97.6%	60.4%	221,578	85.3%	
≥ 4	53,37	0,38	62.8%	98.8%	99.3%	51.2%	243,073	80.8%	
≥ 5	0	0,51	48.8%	100%	100%	43.6%	271,113	74.4%	*
≥ 6	0	0,62	38.1%	100%	100%	39.0%	295,556	69.1%	*
≥ 7	0	0,74	25.6%	100%	100%	34.7%	320,31	62.8%	*
≥ 8	0	0,81	19.1%	100%	100%	32.8%	331,835	59.5%	*
≥ 9	0	0,92	8.4%	100%	100%	30.1%	349,203	54.2%	*
= 10	0	0,98	1.9%	100%	100%	28.7%	358,958	50.9%	*
Univariate Model 2²									
AIRQ score cut-off									
≥ 1	1.44	0	100%	30.4%	48.6%	100%	343,442	65.2%	*
≥ 2	2,05	0,03	98.3%	51.9%	57.4%	97.9%	301,819	75.1%	
≥ 3	2,99	0,11	92.4%	69.1%	66.3%	93.3%	282,227	80.7%	
≥ 4	4,56	0,18	85.7%	81.2%	75.0%	89.6%	266,195	83.5%	
≥ 5	8,46	0,28	74.8%	91.2%	84.8%	84.6%	261,068	83.0%	
≥ 6	16,30	0,38	63.0%	96.1%	91.5%	79.8%	271,116	79.6%	
≥ 7	40,31	0,56	44.5%	98.9%	96.4%	73.1%	306,681	71.7%	
≥ 8	60,84	0,67	33.6%	99.4%	97.6%	69.5%	332,002	66.5%	
≥ 9	0	0,85	15.1%	100%	100%	64.2%	371,925	57.6%	*
= 10	0	0,97	3.4%	100%	100%	61.1%	399,502	51.7%	*

*Quasi-complete separation of data points. The maximum likelihood estimate does not exist at this point. The validity of the model fit is questionable for this cut-off point

¹Model 1 compared well-controlled vs. not well-controlled/very poorly controlled asthma

²Model 2 compared well-controlled/not well-controlled vs. very poorly controlled asthma

³Multivariate-adjusted model for age, sex, and asthma medication

Table 4S. Performance characteristics of the AIRQ questionnaire for the Spanish cohort based on the participant's ACQ-6 score plus exacerbation history

AIRQ Models	Full Analysis Set (FAS)								Model Fit
	LR+	LR-	Sensitivity	Specificity	Positive Predictive Value	Negative Predictive Value	AIC	AUC	
Univariate Model 1¹									
AIRQ score cut-off									
≥ 1	2,17	0,04	97.6%	54.9%	83.3%	90.9%	258,821	76.3%	
≥ 2	4,50	0,14	89.0%	80.2%	91.2%	76.0%	231,477	84.6%	
≥ 3	8,60	0,27	75.6%	91.2%	95.2%	61.9%	246,176	83.4%	
≥ 4	14,37	0,39	63.2%	95.6%	97.1%	53.0%	266,834	79.4%	
≥ 5	22,42	0,52	49.3%	97.8%	98.1%	45.6%	292,648	73.5%	
≥ 6	35,27	0,62	38.8%	98.9%	98.8%	41.3%	310,356	68.8%	
≥ 7	23,51	0,75	25.8%	98.9%	98.2%	36.7%	336,185	62.4%	
≥ 8	17,42	0,82	19.1%	98.9%	97.6%	34.7%	347,968	59.0%	*
≥ 9	0	0,91	8.6%	100%	100%	32.3%	358,691	54.3%	*
= 10	0	0,98	1.9%	100%	100%	30.7%	369,282	51.0%	*
Univariate Model 2²									
AIRQ score cut-off									
≥ 1	1,49	0,02	99.3%	33.3%	55.9%	98.2%	350,198	66.3%	
≥ 2	2,30	0,04	97.8%	57.4%	66.2%	96.9%	291,762	77.6%	
≥ 3	3,70	0,12	91.3%	75.3%	75.9%	91.0%	268,129	83.3%	
≥ 4	5,77	0,21	81.9%	85.8%	83.1%	84.8%	267,648	83.8%	
≥ 5	12,52	0,32	69.6%	94.4%	91.4%	78.5%	268,618	82.0%	
≥ 6	22,89	0,45	56.5%	97.5%	95.1%	72.5%	292,503	77.0%	
≥ 7	63,39	0,61	39.1%	99.4%	98.2%	65.7%	329,023	69.3%	
≥ 8	46,96	0,71	29.0%	99.4%	97.6%	62.2%	356,974	64.2%	
≥ 9	0	0,87	13.0%	100%	100%	57.4%	388,656	56.5%	*
= 10	0	0,97	2.9%	100%	100%	54.7%	411,691	51.4%	*

*Quasi-complete separation of data points. The maximum likelihood estimate does not exist at this point. The validity of the model fit is questionable for this cut-off point

¹Model 1 compared well-controlled vs. not well-controlled/very poorly controlled asthma

²Model 2 compared well-controlled/not well-controlled vs. very poorly controlled asthma

³Multivariate-adjusted model for age, sex, and asthma medication

Table 5S. Agreement between AIRQ, ACT, and ACQ-6 score and physician assessment of asthma control

	Full Analysis Set (FAS)										
	Physician global assessment of patient asthma control										
	Well controlled			Not well controlled			Poorly controlled			Weighted Kappa ¹	95% CI ²
	N	%	95% CI	N	%	95% CI	N	%	95% CI		
AIRQ score assessment	102			124			74				
Missing	0	0%	0%; 3.6%	0	0%	0%; 2.9%	0	0%	0%; 4.9%	0,62	0.55; 0.68
Well controlled	77	75.5%	66.0%; 83.5%	18	14.5%	8.8%; 22.0%	1	1.4%	0.0%; 7.3%		
Not well controlled	25	24.5%	16.5%; 34.0%	63	50.8%	41.7%; 59.9%	11	14.9%	7.7%; 25.0%		
Very poorly controlled	0	0%	0%; 3.6%	43	34.7%	26.4%; 43.7%	62	83.8%	73.4%; 91.3%		
ACT assessment	102			124			74				
Missing	0	0%	0%; 3.6%	0	0%	0%; 2.9%	0	0%	0%; 4.9%	0,71	0.64; 0.77
Well controlled	87	85.3%	76.9%; 91.5%	13	10.5%	5.7%; 17.3%	2	2.7%	0.3%; 9.4%		
Not well controlled	15	14.7%	8.5%; 23.1%	76	61.3%	52.1%; 69.9%	8	10.8%	4.8%; 20.2%		
Very poorly controlled	0	0%	0%; 3.6%	35	28.2%	20.5%; 37.0%	64	86.5%	76.5%; 93.3%		
ACQ-6 assessment	102			124			74				
Missing	0	0%	0%; 3.6%	0	0%	0%; 2.9%	0	0%	0%; 4.9%	0,59	0.51; 0.65
Well controlled	87	85.3%	76.9%; 91.5%	24	19.4%	12.8%; 27.4%	5	6.8%	2.2%; 15.1%		
Not well controlled	12	11.8%	6.2%; 19.6%	45	36.3%	27.8%; 45.4%	6	8.1%	3.0%; 16.8%		
Very poorly controlled	3	2.9%	0.6%; 8.4%	55	44.4%	35.4%; 53.5%	63	85.1%	75.0%; 92.3%		

¹Weighted Kappa was calculated using Cicchetti-Allison weights (linear weights)

²Kappa 1,000 bootstrap sample was used to calculate 95%CI