

SUPPLEMENTARY MATERIAL

SUPPLEMENTARY TABLE 1:

Supplementary Table 1. Selected genetic studies.

Reference	Study type	Population/ Country	Objective	Sample Size	Source	Genes	SNP/Mutation	Results/conclusion		
Adappa et al. 2016 [109]	CG	USA	To determine whether <i>TAS2R38</i> genetics predicts outcomes in CRS patients following sinus surgery	82 CRSwNP 41 CRSsNP	NP, sinonasal tissue	<i>TAS2R38</i>	rs713598 (G/C; Ala/Pro) rs1726866 (G/A; Val-Ala) rs10246939 (T/C; Ile-Val)	The genotype PAV/PAV was related to lower incidence of failing therapy and less frequent sinus surgeries		
Ahmed et al. 2017 [33]	CG	Iraq	To clarify the role of <i>IL4</i> polymorphism in NP	22 healthy controls (HC) 36 NP	NP, inferior turbinate mucosa (ITM)	<i>IL4</i>	?	The polymorphism was found in NP patients but not in controls		
Akygit et al. 2017 [53]	CG	Turkey	To identify genetic polymorphism of <i>SOD2</i> , <i>CAT</i> , <i>iNOS</i> enzymes in E-CRSwNP and NE-CRSwNP patients.	188 HC	Blood	<i>NOS2</i>	-277A/G	The GG genotype (<i>NOS2</i>) and TT genotype (<i>CAT</i>) distributions were different between E-CRSwNP and controls		
				65 E-CRSwNP		<i>SOD2</i>	16C/T			
				65 NE-CRSwNP		<i>CAT</i>	-21A/T			
Alromaih et al. 2013 [27]	pGWAS	Canada	To identify whether genetic factors associated with MHC1 deficiency are present in CRS	196 HC	Blood	<i>CD8A</i>	rs3810831	The minor allele C in <i>CD8A</i> (OR 0.706; p=0.047) and heterozygous CT (OR 0.370; p=0.012) had a protective effect on the development of CRS. The minor allele T in <i>TAPBP</i> (OR 1.53; p=0.009) and heterozygous TT (OR 2.67; p=0.042) were associated with an increased risk for CRS.		
				154 CRSwNP		<i>TAPBP</i>	rs2282851			
				52 CRSsNP						
Al-Shemari et al. 2008 [56]	CG	Canada	To evaluate the effects of SNPs on CRS in a panel of genes related to cysteinyl leukotriene metabolism	200 HC	Blood	<i>ALOX5 AP</i>	rs12430915 rs9506352 rs4769870 rs9579648 rs4076128 rs9579649 rs11616333 rs9315051 rs4769055 rs4420371 rs9578196 rs4466940 rs4293222 rs9578200 rs12429692 rs9285076 rs10162089 rs9670198 rs4254165 rs4319601 rs4356336 rs4769063 rs17612127 rs4238139	Three SNPs located within the <i>ALOX5</i> (rs3780894), <i>CYSLTR1</i> (rs321090) and <i>ALOX5AP</i> (rs17612127) genes reached the nominal p-value threshold (p < 0.05) for association with CRS. However, none of these SNPs resisted multiple testing adjustment.		
				179 CRSwNP			<i>ALOX5</i>		rs3824612 rs12264801 rs3780894 rs3780901 rs2279435 rs7099684 rs1565096 rs1487562 rs7919239 rs2291427 rs7393696 rs2115819 rs11239523 rs4948672 rs7089063	
				27 CRSsNP					<i>CYSLTR2</i>	rs2406939 rs9595961 rs11617224 rs17072059 rs6420296 rs7330127 rs2407249 rs7335898 rs9568087 rs9285169 rs12184704
										<i>CYSLTR1</i>
									<i>LTC4S</i>	
Bae et al. 2013 [110]	CG	Korea	To investigate the association between <i>CIITA</i> and NP	All asthma patients: 158 CRSwNP 309 CRSsNP	Blood	<i>CIITA</i>	rs12932187 rs4781019 rs6498126 rs4781011 rs11074938 rs11074934 rs11074939 rs8043545 rs7404786 rs7201430 rs8063850 rs4781024 rs1139564 rs6498119 rs7189406 rs4774 rs6498124 rs4781016	Two SNPs (rs12932187 and rs11074938) and 2 haplotypes (<i>CIITA</i> _BL1_ht2 and <i>CIITA</i> _BL1_ht5) were demonstrated to be associated with nasal polyps (P=0.001-0.01, OR=0.53-2.35 depending on the genetic model). After multiple testing correction only rs12932187 retained the association with nasal polyps (Pcorr=0.02).		

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Baldan et al. 2015 [111]		CG	Italy	To investigate the effect of 3 <i>IFRD1</i> SPNs on the development of NP in CF patients	CF patients: 40 with NP 103 without NP	Blood	<i>IFRD1</i>	rs6968084 (C>T) rs3807213 (A>C) rs7817 (C>T)	rs7817-CT showed 4-fold higher probability of NP than CC; the TT showed 7.3-fold increased probability. The CAT haplotype showed higher probability of NP (OR 2.63, p=0.004) compared to the CCC haplotype.
Batikhan et al. 2010 [47]		CG	Turkey	To investigate the possible association of <i>TNF</i> -308G/A with NP	95 HC 97 NP patients	Blood	<i>TNF</i>	rs1800629	The presence of the <i>TNF</i> -308 G/A SNP was an independent risk factor for development of NP (OR, 3.68; CI, 1.27–10.7; p = 0.016)
Benito-Pescador et al. 2012 [57]		CG	Spain	To analyze polymorphisms in <i>LTC4S</i> , <i>CYSLTR1</i> , <i>PTGDR</i> , and <i>NOS2</i> as representative genes of inflammation pathways in a population of patients with NP	245 HC 241 NP: 145 asthma 81 NSAID 75 aspirin triad	Blood	<i>LTC4S</i>	rs730012 (-444A>C)	The -444A>C <i>LTC4S</i> polymorphism was significantly associated with NP and atopy (P=.033) and with NP and atopic asthma, (P=.012). A significant association was found when the (CCTTT) repetition of the <i>NOS2A</i> gene was present more than 14 times in patients with NP and asthma (P=.034), in patients with polyposis and intolerance to nonsteroidal anti-inflammatory drugs (P=.009), and in patients with the aspirin triad (P=.005). The <i>PTGDR</i> diplotype CCCT/CCCC (-613CC, -549CC, -441CC and -197TC) was more frequent in patients with NP (P=.043), NP with asthma (P=.013), and the aspirin triad (P=.041)
	<i>CYSLTR1</i>						927 T>C		
	<i>PTGDR</i>						-613 C>T -549 T>C -441 C>T -197 T>C		
	<i>NOS2</i>						CCTTT		
Berghea et al. 2014 [51]		CG	Romania	To investigate the association between <i>TNF</i> SNP with NP in Romanian patients with asthma	45 NP (38 NSAID+ 7 ATA) 61 without NP (8 NSAID+ 53 ATA)	Blood	<i>TNF</i>	rs1799724 (-857 C>T) rs1800629 (-308 G>A) rs361525 (-238 G>A)	There was an association of -857C>T with NP (p=0.01 ATA; p=0.05 NSAID). The allele T was more frequent in NP patients than in non-NP patients.
Bernstein et al. 2009 [30]		CG	USA	To investigate the role of 7 proinflammatory, 4 anti-inflammatory, one Toll receptor and 2 chemokine polymorphism in patients with massive NP	153 HC 179 NP	Buccal cells	<i>TNF</i>	rs1800629	The frequency of the A allele in <i>TNF</i> is significantly higher in patients with NP versus controls (OR 1.86; 95% CI, 1.4–3.09)
	<i>IL1A</i>						rs3783521 rs17561		
	<i>IL1B</i>						rs3087258 rs1143634		
	<i>IL6</i>						rs13447445		
	<i>TGFB1</i>						rs11466315		
	<i>IL10</i>						rs1800895 rs1800894 rs1800896		
	<i>CD14</i>						rs2569190		
	<i>CCL5/ RANTES</i>						rs2107538		
	<i>CCL2</i>						rs3917882		
	Bohman et al. 2017 [21]							GWAS	
<i>HLA-DRA</i>									
<i>BICD2</i>									
<i>VSIR</i>									
<i>SLC5A1</i>									
Bosse et al. 2009 [73]		GWAS	Canada	To perform pooling-based GWAS in two case-control cohorts, one of them consisting of patients with CRSwNP	189 HC 210 with severe CRS 157 CRSwNP 53 CRSsNP	Blood	<i>LAMA2</i>	rs2571584	600 SNPs from 445 genes that were potentially associated with CRS (P < 0.05). Each of these novel high-priority SNPs had allele frequency differences between cases and controls at a level worthy of additional investigation. The most significant SNP for each of the top 10 genes are shown in this table.
	<i>PARS2</i>						rs2873551		
	<i>NAV3</i>						rs1726427		
	<i>LAMB1</i>						rs4727695		

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							<i>CACNA1I</i>	rs3788568			
							<i>KIAA1456</i>	rs11779957			
							<i>MUSK</i>	rs10817091			
							<i>TRIP12</i>	rs10535833			
							<i>AOAH</i>	rs4504543			
							<i>MSRA</i>	rs7001821			
Bussu et al. 2007 [69]		CG	Italy	To evaluate the potential involvement of <i>ADRB2</i> A16G polymorphism in sinonasal polyposis	47 HC 56 NP	Blood	<i>ADRB2</i>	rs1042713 (g.5285A>G)	The presence of Arg (A allele) is significantly higher in NP patients than in controls (p=0.0386)		
Buysschaert et al. 2010 [41]		GWAS	Belgium	To investigate whether certain SNPs predispose to NP	415 HC 273 NP	Blood	<i>IL1RL1</i>	rs1420101	Rs3939286 was significantly associated with NP (OR 1.60; 95% CI 1.16-2.22; p=0.0041). The A-allele conferred a risk for NP (OR 1.53; 95% CI 1.21-1.96; p=0.0041). Rs1420101 may increase risk when in combination with rs3939286		
							<i>IKZF2</i>	rs12619285			
							<i>GATA2</i>	rs4431128			
							<i>IL5</i>	rs4143832			
							<i>SH2B3</i>	rs3184504			
							<i>WDR36</i>	rs2416257			
							<i>MHC</i>	rs2269426			
							<i>MYB</i>	rs9494145			
							<i>GFRA2</i>	rs748065			
						<i>IL33</i>	rs3939286				
Cantone et al. 2018 [61]		CG	Italy	To investigate the relevance of <i>TAS2R38</i> genetic variants in the susceptibility to bacterial infections	100 CRSwNP	Saliva, blood	<i>TAS2R38</i>	rs713598 (C145G; Pro>Ala) rs1726866 (C785T; Ala>Val) rs10246939 (G886A;Val>Ile)	The nonfunctional genotype (AVI) is more frequent among CRS patients than in the general population (25% vs. 18.4%, P=0.034). No relationship with severity was found.		
Castano et al. 2009 [36]		CG	Canada	To investigate whether certain polymorphisms in the <i>IL1RL1</i> gene are differentially present in patients with surgery-unresponsive CRS and in control subjects	187 HC 154 CRSwNP 52 CRSsNP	Blood	<i>IL1RL1</i>	rs974389	rs10204137	Statistically significant allelic associations with CRS were noted for 5 SNPs (rs10204137, p=0.04; rs10208293, p=0.03; rs13431828, p=0.008; rs2160203, p=0.03, and rs4988957, p=0.03). But only one SNP significantly associated with CRSwNP (rs13431828, p=0.008)	
								rs985523	rs10208293		
								rs1041973	rs12105808		
								rs1420103	rs12712142		
								rs1921622	rs12996097		
rs2160203	rs13431828										
							rs3771177	rs17696274			
							rs4988957				
Castano et al. 2010 [71]		CG	Canada	To assess the association of polymorphisms in the <i>MET</i> gene with CRS in a Canadian population	196 HC 154 CRSwNP 52 CRSsNP	Blood	<i>MET</i>	rs38840	rs2237711	rs1024658	The genotype distribution of two SNPs in the <i>MET</i> gene (rs388840, rs38850) displayed a significantly different genotypic distribution (p≤ 0.05) between CRS patients and controls. The most significant association in the <i>MET</i> gene was found with SNP rs38850 (p=0.004).
								rs10271561	rs10243024		
								rs40239			
								rs714180	rs38855		
								rs38841	rs38857		
								rs39747	rs2237717		
								rs38845			
								rs38846	rs2299440	rs1752698	
								rs7798983	rs2402118		
								rs38849	rs193688		
							rs722134	rs1621			
							rs38850	rs42336			
Cheng et al. 2006 [34]		CG	Taiwan	To assess the association of <i>IL1B</i> and <i>IL1RN</i> gene polymorphisms with CRS	103 HC 88 CRS 61 CRSwNP 27 CRSsNP	Blood	<i>IL1B</i>	-511C>T +3953C>T	There were significant differences in the distribution of the <i>IL1RN</i> polymorphism between the control subjects and patients with CRS (P=.05). The II allele of <i>IL1RN</i> occurred more frequently in the CRS patient		
							<i>IL1RN</i>	Variable number tandem repeat of an 86–base pair segment in intron 2			

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										group (OR 3.3; 95% CI, 1.25-9.18, P=0.01).
Cormier et al. 2009 [112]		CG	Canada	To determine whether SNP in <i>TNF</i> , <i>TNFAIP3</i> , and <i>TNFAIP6</i> genes were associated with CRS	196 HC 206 CRS 154 CRSwNP	Blood	<i>TNF</i>	rs2229094 rs1121800 rs1321136r s1800750 rs2256965 rs2256974 rs2857706 rs3093561	rs3093672 rs769177 rs77669888 rs9267502 rs9469027 rs1800629 rs361525 rs4987027	Two polymorphisms in <i>TNFAIP3</i> (rs3757173 and rs5029938) are weakly associated with severe CRS but no association was found with genetic variants in <i>TNF</i> or <i>TNFAIP6</i> . None was associated to risk of NP.
							<i>TNFAIP3</i>	rs5029963 rs5029935 rs5029939 rs5029965	rs3757173 rs5029938 rs661561 rs610604	
							<i>TNFAIP6</i>	rs6433371 rs12466578 rs2342910 rs3771889 rs3771891 rs10432475	rs16830015 rs6707824 rs10183099	
Dar et al. 2018 [65]		CG	India	To assess the risk of CRSwNP conferred by SNPs in <i>FcεR1α</i> gene in a North Indian cohort	50 HC 100 CRSwNP	Blood	<i>FCER1A</i>	rs2427827 rs2251746 rs2298804 rs2298805 rs2269718		In those cases with high serum IgE, the T allele of rs2427827 is significantly more frequent in CRSwNP patients (OR 2.24; p=0.02)
De Alarcon et al. 2016 [58]		CG	USA	To evaluate the association of <i>LTC4S</i> and <i>PAI-1</i> variants with CRS	66 HC 16 CRSsNP 117 CRSwNP	Blood, polyp fibroblasts	<i>LTC4S</i> <i>SERPINE1 (PAI1)</i>	rs730012 (-444A>C) rs1799762 (4G/5G ins.)		The C allele of <i>LTC4S</i> was more frequent in those NP patients also diagnosed with chronic hyperplastic eosinophilic sinusitis (p<0.04)
Ekinci et al. 2011 [70]		CG	Turkey	To examine whether there is an association of eotaxin-1 (<i>CCL11</i>) gene polymorphisms with NP	93 HC 85 NP	Blood	<i>CCL11</i>	rs1490392522 (-384A>G) rs762429865 (67 G>A)		The selected SNPs are more frequent in NP patients than in HC (p=0.044 and p=0.019, respectively). However, their relation was statistically poor (association coefficient =0.18).
Erbek et al. 2007 [35]		CG	Turkey	To investigate the association between NP and SMPs of the proinflammatory cytokines <i>IL1A</i> , <i>IL1B</i> , <i>TNFα</i> .	106 HC 82 NP	Blood	<i>IL1A</i> <i>IL1B</i> <i>TNF</i>	4845G>T -511C>T rs361525 (-238 G>A) rs 1800629 (-308G>A)		The 4845G and 4845TT genotypes of the <i>IL1A</i> gene were associated with NP (P<.05). The frequency of the <i>IL1B</i> -511 CC and CT were significantly higher in patients with NP than in controls (P=.01). There was a significantly high risk of susceptibility to NP in patients with the -308 GA genotype (P=.001).
Esmailzadeh et al. 2015 [19]		CG	Iran	To investigate the association of <i>HLA-DRB</i> and <i>-DQ</i> genetic variabilities in patients with AERD	100 HC 50 CRSwNP + asthma	Blood	<i>HLA-DRB1</i>	<i>HLA-DRB1*0101</i> <i>HLA-DRB1*15</i> <i>HLA-DRB1*16</i> <i>HLA-DRB1*0301</i> <i>HLA-DRB1*04</i> <i>HLA-DRB1*07</i> <i>HLA-DRB1*08</i> <i>HLA-DRB1*0901</i> <i>HLA-DRB1*1001</i> <i>HLA-DRB1*11</i> <i>HLA-DRB1*12</i> <i>HLA-DRB1*1301</i> <i>HLA-DRB1*1302</i>		Two variations are associated with increased risk of NP: <i>HLA-DRB4</i> , OR 2.34, CI 1.37–4.00, p>0.01 <i>HLA-DQB1*0302</i> , OR 4.56, CI 2.10–9.91, p<0.01 Two variations are associated with reduced risk of NP: <i>HLA-DRB3</i> , OR 0.41, CI 0.24–0.68, p<0.01 <i>HLA-DQB1*0301</i> , OR 0.39, CI 0.21–0.73, p<0.01

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								<p><i>HLA-DRB1*1305</i></p> <p><i>HLA-DRB1*14</i></p>	
							<i>HLA-DRB3</i>	<i>HLA-DRB3</i>	
							<i>HLA-DRB4</i>	<i>HLA-DRB4</i>	
							<i>HLA-DRB5</i>	<i>HLA-DRB5</i>	
							<i>HLA-DQA1</i>	<p><i>HLA-DQA1*0101</i></p> <p><i>HLA-DQA1*0102</i></p> <p><i>HLA-DQA1*0103</i></p> <p><i>HLA-DQA1*0104</i></p> <p><i>HLA-DQA1*0201</i></p> <p><i>HLA-DQA1*0301</i></p> <p><i>HLA-DQA1*0401</i></p> <p><i>HLA-DQA1*0501</i></p>	
							<i>HLA-DQB1</i>	<p><i>HLA-DQB1*0201</i></p> <p><i>HLA-DQB1*0301</i></p> <p><i>HLA-DQB1*0302</i></p> <p><i>HLA-DQB1*0303</i></p> <p><i>HLA-DQB1*0305</i></p> <p><i>HLA-DQB1*0401</i></p> <p><i>HLA-DQB1*0501</i></p> <p><i>HLA-DQB1*0601</i></p> <p><i>HLA-DQB1*0602</i></p> <p><i>HLA-DQB1*0604</i></p>	
Fajardo-Dolci et al. 2006 [20]		CG	Mexico	To determine the contribution of the human major histocompatibility complex <i>HLA-DQA1</i> , – <i>DQB1</i> , and <i>TNFα</i> genes with simple nasal polyposis.	151 HC 31 NP	Blood	<i>HLA-DQA1</i>	<p><i>HLA-DQA1*0101/4</i></p> <p><i>HLA-DQA1*0102</i></p> <p><i>HLA-DQA1*0103</i></p> <p><i>HLA-DQA1*0201</i></p> <p><i>HLA-DQA1*030101</i></p> <p><i>HLA-DQA1*0401</i></p> <p><i>HLA-DQA1*0501</i></p>	<p>The allele <i>HLA-DQA1*0201</i> was found to be involved in susceptibility to develop simple NP, without asthma, aspirin intolerance, or any other allergic diseases. OR 6.79 CI (1.9-23.9)</p> <p>13% etiological fraction was found for the haplotype <i>HLA-DQA1*0201-DQB1*0201</i> (P=0.016).</p>
							<i>HLA-DQB1</i>	<p><i>HLA-DQB1*0201</i></p> <p><i>HLA-DQB1*0301</i></p> <p><i>HLA-DQB1*0302</i></p> <p><i>HLA-DQB1*0402</i></p> <p><i>HLA-DQB1*0501</i></p> <p><i>HLA-DQB1*0502</i></p> <p><i>HLA-DQB1*0503</i></p> <p><i>HLA-DQB1*0504</i></p> <p><i>HLA-DQB1*0601</i></p>	

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								HLA-DQB1*0602 HLA-DQB1*0603	
							TNF	rs1800629 (-308 G>A) rs361525 (-238 G>A)	
Fruth et al. 2011 [113]		CG	Germany	To analyze the potential association of <i>GST</i> polymorphisms and CRS.	52 HC 118 CRS 69 CRSwNP 49 CRSsNP	NP, ITM	<i>GST</i>	GST-T1 GST-M1 GST-P1	No correlation
Fruth et al. 2012 [114]		GC	Germany	To shed light on the significance of <i>SPINK5</i> and the development of inflammatory diseases of the upper respiratory tract.	30 HC 59 CRSwNP 15 CRSsNP	NP, ITM	<i>SPINK5</i>	rs17775319 (G1258A) G2475T rs1243172589 (A2915G) rs745601984 (A1103G)	No correlation
Gallo et al. 2016 [62]		CG	Italy	To confirm the proposed correlation between <i>TAS2R38</i> genotype, CRS, and related comorbidities.	39 HC 36 CRSwNP 17 CRSsNP	Blood	<i>TAS2R38</i>	rs713598 (G/C; Ala/Pro) rs1726866 (G/A; Val-Ala) rs10246939 (T/C; Ile-Val)	No differences found in genotypic distribution
Henmyr et al. 2014 [37]		GWAS	Turkey, Finland, China, Korea, USA, Belgium	To investigate the reproducibility of previous SNP associations with CRSsNP and CRSwNP.	1588 HC from Illumina data base 613 Belgian patients: 275 CRSwNP 338 CRSsNP	Blood, database	<i>IL1A</i> <i>IL1B</i> <i>RYBP</i> <i>DCBLD2</i> <i>TNFA</i> <i>AOAH</i> <i>IL33</i> <i>IRAK4</i> <i>IL10</i> <i>CACNG6</i> <i>MMP9</i>	rs17561 rs16944 rs4532099 rs828618 rs1800629 rs361525 rs4504543 rs3939286 rs4251431 rs1800870 rs192808 rs3918242 rs17577	Some SNPs are associated with increased risk of NP: <i>IL1A</i> rs17561 <i>RYBP</i> rs4532099 <i>TNF</i> rs1800629 <i>IL33</i> rs3939286 <i>IL10</i> rs1800870 <i>CACNG6</i> rs192808 <i>MMP9</i> rs3918242 Some SNPs are associated with reduced risk of NP <i>IL1B</i> rs16944 <i>DCBLD2</i> rs828618 <i>AOAH</i> rs4504543 <i>IRAK4</i> rs4251431
Henmyr et al. 2016 [115]		CG	Sweden, 1000Genomes Project	To screen for rare variants in <i>PARS2</i> and to evaluate for accumulation of such variants in CRS patients.	372 HC 138 CRSwNP 172 CRSsNP	Blood	<i>PARS2</i>	rs143717155 rs35201073 rs370234936 rs2270004 rs145005088 rs1180946 rs11577368 rs145866387 rs74617964 rs1180947 rs116816976 rs12023572 rs768053281 rs61768813 rs563439229 rs1180945 rs116416055	A significant surplus of variation was observed in the CRS patients (p=0.048). Haplotype analysis of the region showed a significant excess of rare haplotypes in the CRS patients compared to HC in the following SNPs: rs2873551, rs2270004, rs11577368, rs1180946, rs1180945 TTAGC p=0.0048 TTCCC p=0.0048 TCAGT p=0.0016
Hytönen et al. 2001 [67]		CG	Finland	To investigate if the frequency of the most common <i>CFTR</i> mutations was more	135 CRS	Blood	<i>CFTR</i>	ΔF508	No abnormal distribution was observed in CFR patients

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				common among CRS patients with or without NP.	91 CRSsNP 46 CRSwNP			394delTT		
Ismi et al. 2017 [50]		CG	Turkey	To determine the genetic susceptibility of NP formation to TNF and IL1B polymorphisms	91 HC 71 CRSwNP	Blood	TNF	-308G>A	There was a statistically significant increase in the expression of the TNF -308 GG and IL1B -511 CC genotypes in the patients with NP	
							IL1B	-511T>C		
Karjalainen et al. 2003 [38]		CG	Finland	To establish whether IL1A and IL1B have an effect on susceptibility to NP.	35 CRSwNP 210 CRSsNP	Blood	IL1A	4845G>T	The risk of NP was markedly increased in IL1A allele G homozygous subjects (OR 2.73; 95%CI 1.40–5.32, p=0.005). In the case of IL1B no significant associations were found.	
							IL1B	-511C>T		
Keles et al. 2008 [22]		CG	Turkey	To evaluate whether there is a relationship between HLA-A, -B, -Cw, and -DRB1 alleles and developing NP.	100 HC 66 NP	Blood	HLA-A	HLA-A *01 HLA-A *02 HLA-A *03 HLA-A *11 HLA-A *24 HLA-A *26 HLA-A *33	HLA-B*07 and -Cw*12 alleles were significantly higher in the NP patients than in the control group. HLA-B*57 and -Cw*04 alleles were significantly lower in the NP patients than in the control group. In the NP patients with ASA, there was a significant increased frequency of the HLA-A*24, -Cw*01, -Cw*12, and -DRB1*04 alleles. HLA-A*33 and -Cw*12 alleles in NP patients who had polypectomy history were significantly higher than in the control group. In NP patients, a significantly decreased frequency of the HLA-Cw*04 and -DRB1*11 alleles was shown.	
							HLA-B	HLA-B *07 HLA-B *15 HLA-B *35 HLA-B *44 HLA-B *51 HLA-B *57		
							HLA-Cw	HLA-Cw *01 HLA-Cw *02 HLA-Cw *03 HLA-Cw *04 HLA-Cw *06 HLA-Cw *07 HLA-Cw *08 HLA-Cw *12		
							HLA-DRB1	HLA-DRB1*01 HLA-DRB1*03 HLA-DRB1*04 HLA-DRB1*07 HLA-DRB1*11 HLA-DRB1*14		
Kilty et al. 2010 [116]		CG	Canada	To investigate the association between SNPs in the SERPINA1 gene and CRS	196 HC 154 CRSwNP 27 CRSsNP	Blood	SERPINA1	rs11558262 rs11832 rs1243160 rs1243163 rs1243166 rs1243167 rs1243168 rs1243169 rs1243171 rs12884390 rs1303 rs17287271 rs17580	rs1956707 rs2071274 rs2230075 rs2239651 rs2749531 rs2753934 rs3748316 rs4900229 rs4900230	Two SNPs (rs1243168 and rs4900229) were associated with the disease. rs1243168 T allele was significantly associated with severity (p<0.01)

Supplementary Table 1. Selected genetic studies.

								rs17751614 rs17751769 rs17824797	rs4905198 rs6575424 rs6647 rs709932 rs7151526 rs8010121 rs877081	
Kim et al. 2007 [117]		CG	Korea	To evaluate the association of <i>TGFB1</i> polymorphisms with an AERD phenotype in the Korean population	456 HC 206 AERD 72 NP 324 ATA 10 NP	Blood	<i>TGFB1</i>	rs13447445		No association with NP
Kim et al. 2009 [64]		CG	Korea	To evaluate associations between genetic polymorphisms of adenosine deaminase and adenosine receptors with the AERD phenotype	183 HC 136 AERD 47 NP 181 ATA 10 NP	Blood	<i>ADA</i>	rs11086932 rs244076	No association with NP was described. Significant differences between normal and patients with AERD in the ADORA1 SNP genotype frequencies for rs16851030 (P=0.001) and rs6664108 (P=0.013).	
	<i>ADORA1</i>						rs10920568 rs6664108 rs6427994 rs16851030 rs12744240			
	<i>ADORA2A</i>						rs5996696 rs5751876			
	<i>ADORA3</i>						rs2298191 rs10776727 rs1544224 rs2229155			
Kim et al. 2012 [23]		CG	Korea	To investigate the associations of HLA-DRA polymorphisms with NP in asthmatic and AERD patients.	158 CRSwNP 309 CRSsNP	Blood	<i>HLA-DRA</i>	rs9268628 A>C rs3129871 C>A rs9268633 G>A rs9405035 G>A rs14004 C>A rs9268644 C>A rs9268645 G>C rs3129878 A>C rs3135392 G>T rs6926374 G>A rs3129881 C>T rs17496549 C>T rs6911777 T>C rs3129886 C>T rs8084 C>A rs2239805 A>C rs2239804 G>A rs7192 G>T rs4935354 A>G rs7194 A>G	4 SNPs were significantly associated with NP Rs9268644 p=0.009 Rs3129878 p=0.033 Rs3129881 p=0.013 Rs2239805 p=0.029 And the haplotype (rs3129871; rs8084; rs2239805; rs2239804; rs7192; rs4935354; rs7194; rs1051336; rs1041885) TAAATGGA (p=0.029)	

Supplementary Table 1. Selected genetic studies.

								rs1051336 G>A rs1041885 A>T	
Kostuch et al. 2005 [66]		CG	Poland	To determine the prevalence of the most common CFTR mutations in patients with NP without suspicion of CF.	70 human placentas 44 NP	Blood, placenta	CFTR	ΔF508 G551D G542X N1303K 1717-1G>A W1282X R553X ΔI507	ΔF508 is more frequent in patients than in HC (p=0.0032) and in the general Polish population as well (P =0.0059).
Kosugi et al. 2013 [118]		CG	Brazil	To analyze the relationship between an IL6 polymorphism and asthmatic NP patients.	81 HC 45 asthmatic with NP 63 non asthmatic NP 45 asthmatic without NP	Blood	IL6	rs374295772 (-174G>C)	Genotype distribution was non-significant, but GG genotype appeared more frequently in all inflammatory groups.
Kristjánsson et al. 2019 [55]		GWAS	Iceland, UK	To search for sequence variants affecting the risk of NP or CRS	Iceland 353939 HC 3188 cases UK 406147 HC 2420 cases	Database	HLA-DQA1	rs1391371	The mentioned variants at ten loci were found that associate with NP at genome-wide significance. The variant with the largest effects on the risk of NP is a low-frequency missense variant rs34210653[A] (Thr560Met) in ALOX15 that confers a 68% reduction in NP risk (p= 8.0x10 ⁻²⁷ OR 0.32, 95% CI 0.26, 0.39).
							IL33	rs1888909	
							TSLP	rs1837253	
							ALOX15	rs34210653	
							10p14	rs1444782	
							FOXP1	rs17718444	
							CYP2S1	rs338598	
							IL18R1	rs6543124	
							SLC22A4	rs1050152	
							MYRF	rs174535	
Kuran et al. 2019 [39]		CG	Turkey	To analyze possible genetic factors that increase susceptibility to NP.	98 HC 78 NP	Blood	IL1RN	rs2234663	Distribution of genotypes of IL1RN and IL4 was significantly different in NP vs HC (p=0.0001)
							IL4	rs8179190	
							IL2	rs206976	
Luxenberger et al. 2000 [119]		CG	Austria	To determine the association of HLA-A, -B, -DR, and -DQ with NP	1070 HC 89 NP	Blood	HLA-A HLA-B HLA-DR HLA-DQ	A significant association was seen with HLA-A74 and nasal polyposis	
Mfuna Endam et al. 2009 [29]		CG	Canada	To explore association between SMPs in IL22RA1 and severe CRS	196 HC 206 CRS 154 CRSwNP 52 CRSsNP	Blood	IL22RA1	rs10465895 rs2502450 rs10751768 rs3795300 rs10794665 rs3936073 rs10903031 rs11249201 rs4292900 rs11577442 rs4648936 rs11578307 rs4648942 rs11579657 rs12070496 rs4649187 rs12092673 rs6424157 rs12408946 rs7418238 rs16829225 rs7513249	Three SNPs (rs4292900 Pnom =0.0006, OR 1.757; rs4648936 Pnom=0.0011, OR 1.716; rs16829225 Pnom=0.0014, OR 1.977) show significant differences in allelic frequencies between cases and controls
Mfuna Endam et al. 2010 [28]		CG	Canada	To replicate the CRS associations recorded for IL1A, IL1B, and TNF	196 HC	Blood	IL1A	rs17561 rs3783521 rs1800587 rs3783538 rs2048874	For rs17561, this study replicated previous results about the association of the TT homozygote genotype (OR, 3.39; P=.007). The protective effect of

Supplementary Table 1. Selected genetic studies.

				in a cohort of Canadian patients with severe CRS.	206 CRS 154 CRSwNP 52 CRSsNP			rs2856838 rs6722023 <i>IL1B</i> rs16944 <i>TNF</i> rs1121800 rs13211368 rs1800629 rs1800750 rs2229094 rs2256965 rs2256974 rs2857706 rs3093561 rs3093672 rs361525 rs4987027 rs769177 rs7766988 rs9267502 rs9469027	rs2856838 (OR, 0.38; P=.002) and the risk effect of rs1800587 (OR=3.16, P=.008) are enhanced with the homozygote form of the minor allele. In contrast, no association was found with SNPs in <i>IL1B</i> or <i>TNF</i> .
Mfuna Endam et al. 2014 [59]		GWAS	Canada	To identify taste receptor associated with CRS and verify whether known SNP replicated in their CRS cohort	GCRS1 196 HC 206 CRS GCRS2 190 HC 408 CRSwNP	Blood	<i>TAS2R1</i> rs17788846 rs41483 rs12374524 rs6555620 rs10746553 rs1015855 <i>TAS2R10</i> rs669503 rs10845219 <i>TAS2R13</i> rs1015442 rs1015443 <i>TAS2R14</i> rs3851586 rs1013311 rs3741843 <i>TAS2R3</i> rs765007 rs6962760 <i>TAS2R38</i> rs4726481 rs10246939 rs1726866 <i>TAS2R39</i> rs11979433 <i>TAS2R4</i> rs2234002 rs2190245 <i>TAS2R41</i> rs2966709 rs2966715 rs2949746 rs2949770 rs12536735 rs1806578 rs2966701 rs2966699 <i>TAS2R43</i> rs2966699 rs35911096 rs2708333 rs2597975 rs2708364 rs2599396 rs7313683 rs2597974 <i>TAS2R44</i> rs4763616 rs2010481 <i>TAS2R46</i> rs2708389 rs11533164 rs2708377 rs2255418 <i>TAS2R48</i> rs10772420	57 SNPs in <i>TAS2Rs</i> and 16 SNPs in <i>TAS1Rs</i> had allele frequency differences above 10% between controls and patients (range, 10.2% to 32.4%). Three coding SNPs associated with CRS were identified: 1 in the <i>TAS2R13</i> gene (rs1015443, biallelic differences of 13.8% in GCRS1), and 2 others in the <i>TAS2R49</i> gene (rs12226920, biallelic difference of 16.0% in GCRS1; and rs12226919, biallelic difference of 11.9% in GCRS1)	

Supplementary Table 1. Selected genetic studies.

							<p><i>TAS2R49</i> rs1463237 rs10772408 rs4298989 rs12226920</p> <p>rs12581501 rs12226919 rs11054150</p>	
							<p><i>TAS2R5</i> rs35010424</p> <p>rs11769672</p> <p>rs11773137</p> <p>rs1859646</p>	
							<p><i>TAS2R50</i> rs2900554</p> <p>rs6488331</p>	
							<p><i>TAS2R7</i> rs7313019</p>	
							<p><i>TAS1R1</i> rs11122100</p> <p>rs12080675</p>	
							<p><i>TAS1R2</i> rs28410948 rs6686865 rs7417542 rs9662598</p> <p>rs7411042 rs6685177 rs12137730 rs4920566</p> <p>rs6603912 rs6684577 rs12063142 rs3935570</p> <p>rs12042960 rs7418296</p>	
Molga et al. 2016 [120]		CG	Poland	To assess genetic predisposition of MMP1 -1607 G/GG polymorphism to CRSwNP	463 HC 206 CRSwNP	Blood	<p><i>MMP1</i> rs199750 (-1607 G/GG)</p>	The frequency of genotypes was not significant related to CRSwNP, but GG is relates to increases number of surgeries (p=0.002) and bronchial asthma (p=0.021)
Molnar-Gabor et al. 2000 [24]		CG	Hungary	To investigate whether there is an association between HLA-DRB1, -DQA1, and DQB1 alleles and developing NP	50 HC 50 CRSwNP	Blood	<p><i>HLA-DR5</i></p> <p><i>HLA-DR7</i></p> <p><i>HLA-DQA1</i> HLA-DQA1*0101 HLA-DQA1*0102 HLA-DQA1*0103 HLA-DQA1*0104 HLA-DQA1*0201 HLA-DQA1*0301 HLA-DQA1*05011 HLA-DQA1*05012</p> <p><i>HLA-DRB1</i> 1-10</p> <p><i>HLA-DQB1</i> HLA-DQB1*0201 HLA-DQB1*0202 HLA-DQB1*0302 HLA-DQB1*0301 HLA-DQB1*0303 HLA-DQB1*0402 HLA-DQB1*0501 HLA-DQB1*0502 HLA-DQB1*0503 HLA-DQB1*0601 HLA-DQB1*0602 HLA-DQB1*0603</p>	The odds ratios for developing nasal polyposis increased in people carrying the DR7 (OR 2.55) allele with the linked DQA1*0201 (OR 2.52) and DQB1*0202 (OR 5 2.84) alleles. On the other hand, DR5 (OR 0.66) linked with DQA1*05012 (OR 0.69), DQB1*0301 (OR 0.57) alleles showed a decreased odds ratio value.

Supplementary Table 1. Selected genetic studies.

								HLA-DQB1*0604	
Mrowicka et al. 2014 [32]		CG	Poland	To investigate the relationship between IL1B and IL4 promoter polymorphisms	200 HC 208 CRSwNP	Blood	IL1B	rs55778004 (-511C>T)	The TT genotype for C-511T mutation associated with the risk of developing NP in a Polish population
							IL4	-590C>T	
Nakayama et al. 2020 [121]		GWAS	Japan	To perform an association study of CRSwNP and AERD with genetic variants in the TSLP locus	1908 HC 499 CRSwNP	Blood	TSLP	rs1751303 rs3806933 rs10056340 rs1898671 rs1837253 rs2416257 rs3806932 rs1438673	Significant association between CRSwNP and rs1837253, rs3806932 and rs3806933, with the most significant association being observed at rs1837253 (p= 1.27x10 ⁻⁶ ; OR, 1.60; 95% CI, 1.32-1.94)
Ozdas et al. 2015 [122]		CG	Turkey	To analyze SNPs of the RYD5 gene and to determine the effect on poly p formation	238 HC 196 NP	Blood	RYD5	rs113795008 rs7951297 rs535294582 rs2294083 rs2280540 rs2294082 rs144999256 rs61997072 rs148962288	Four SNPs (rs113795008, rs2280540, rs2294083, and rs2294082) were significantly associated with NP. The individuals with combined genotypes of six risk alleles (rs113795008, rs2280540, rs7951297, rs2294083, and rs2294082) had significantly higher risks for NP compared with the ones with one or four risk alleles.
Palikhe et al. 2017 [123]		CG	Korea	To investigate the potential associations between ABCC4 gene polymorphisms and asthma genotype.	120HC 270 asthma 106 NP	Blood	ABCC4	rs868853 (A>G) rs839951 (C>G)	No significant association
Park et al. 2006 [43]		CG	Korea	To evaluate expression of cyclooxygenase (COX)-2 and 5-lipoxygenase (5-LO) associated with IL4 promoter polymorphism -590 in NP tissues	70 HC 61 NP	Blood	IL4	-590 C>T	A T>C exchange at -590 position was correlated with NP. The T allele was significantly more frequent in NP (p=0.028).
Pasaje et al. 2012a [124]		CG	Korea	To explore the association of DCBLD2 gene with the presence of NP in asthma patients	309 HC 158 NP+asthma	Database	DCBLD2	rs2439224 rs828618 rs1371687 rs828616 rs9838238 rs16840208 rs17278047 rs17270986 rs7615856 rs1062196 rs828621 rs8833	Six SNPs were associated with the presence of NP: rs1371687, rs7615856, rs828621, rs828618, rs828616, and rs8833. After multiple testing adjustment, only rs828621 remained significant (p=0.006)
Pasaje et al. 2012b [125]		CG	Korea	To investigate the association between EMID and NP	309 asthma no NP 158 asthma+NP	Database	EMID	rs6945102 rs1476652 rs4729697 rs6973489 rs10237610 rs7802156 rs9986717 rs10953342 rs10254516 rs12668018 rs10239458 rs1008064 rs221 rs13232646 rs10435333 rs1543883 rs6944691 rs1859625 rs6942770 rs6947089 rs9640666 rs9969331 rs6947185 rs12538381 rs11770876 rs17135512 rs11772022 rs1558015 rs11772003 rs10250055 rs10223928 rs6947735 rs4729705 rs2158739 rs10254310 rs10279545	Ten EMID2 SNPs (rs6945102, rs4729697, rs221, rs10435333, rs6947185, rs4727494, rs13233066, rs1008064, rs1543883, and rs13245946) were associated with the presence of nasal polyps (p= 0.004- 0.05, OR 0.61-1.32) depending on the genetic model. rs6945102, rs4729697, rs221, and rs10435333, were found to be significantly associated with NP in the overall Korean asthma patients even after multiple testing corrections

Supplementary Table 1. Selected genetic studies.

								rs4045 rs6949799 rs4727491 rs13238748 rs4727494 rs13233066 rs869127	rs6945961 rs13245946 rs17470799 rs10237510 rs17135617 rs17135621	
Pascual et al. 2008 [126]		CG	Spain	To analyze the (CCTTT) _n polymorphism of NOS2 in patient with NP and/or asthma	98 HC 46 NP 150 asthma	Blood	NOS2	(CCTTT) _n		Allele frequency distribution is significantly different between HC and NP (p=0.003). A 15-repeat cutoff is associated with increased risk of NP (OR 14.39; 95% CI, 3.02 - 68.60; P = .001)
Pavon-Romero et al. 2018 [54]		SNP array	Mexico	To evaluate whether contribution to susceptibility of SNPs reported in other populations are associated with AERD in Mexican patients	179 HC 120 AERD 179 asthma	Blood	ACE	rs4309† rs4293†	In AERD vs. HC, we identified 22 associated SNPs, with 11 SNPs associated with risk in 7 genes (ACE, MS4A2, FSIP2, IL10, FCER1G, KIFC3, and ANX4; denoted as † in the adjacent column). Two SNPs were strongly associated: ACE rs4309 (C allele p = 0.0001, OR = 1.92, CI 95% = 1.37–2.69) and MS4A2 rs573790 (C allele p = 0.0002, OR = 1.94, CI 95% = 1.35–2.79). By contrast, 11 SNPs in 5 genes (PPARG, IL10, RGS7BP, TBXAS1, and FANCC) were associated with protection.	
	MS4A2						rs576790† rs502581†			
	FSIP2						rs2631700† rs2631702†			
	KIFC3						rs2285700†			
	ANX4						rs7588022†			
	FCER1G						rs4489574† rs7528588			
	IL10						rs1800896† rs3024498† rs1554286 rs1800872			
	PPARG						rs2960421 rs4135275 rs1875796			
	RGS7BP						rs6870654			
	TBXAS1						rs13239058 rs10487667 rs6962291			
							FANCC	rs1326188		
Purnell et al. 2019 [63]		CG	USA	To determine the frequency of 6 SNPs in genes with bitter taste signaling function.	1000 Genomes database 74 CRS 41 CRSwNP 33 CRSsNP	Buccal cells	TAS2R38	rs713598	No differences between CRSwNP and CRSsNP	
	GNB3						rs5443			
	TAS2R19						rs10772420			
	TAS2R20						rs12226920			
	RGS21						rs7528947 rs1175152			
Ramirez-Anguiano et al. 2006 [25]		CG	Mexico	To determine the association of HLA-DRB1 alleles with NP in the Mexican Mestizo population.	99 HC 34 NP	Blood	HLA-DRB1	HLA-DRB1*02 HLA-DRB1*03 HLA-DRB1*04 HLA-DRB1*05 HLA-DRB1*07 HLA-DRB1*08	Significant increase in the *03 and *04 (OR 2.2, p=0.009) allele frequencies. Significant decrease in the *08 allele (OR 0.2, p=0.01)	

Supplementary Table 1. Selected genetic studies.

Sachse et al. 2010 [127]		CG	Germany	To detect staphylococcal colonization in nasal polyps and controls.	51 HC 68 NP	NP, ITM	<i>TLR2</i>	rs5743708	The minor allele A is not associated with NP
Sitarek et al. 2012 [52]		CG	Poland	To investigate the association of COX-2 and MET gene polymorphisms with the risk of CRSwNP.	200 HC 195 NP	Blood	<i>COX2</i>	rs20417	Increased risk ($p > 0.001$) of CRSwNP associated with the C allele of COX2 (OR 6.05) and G allele of MET (OR 5.52) The combined genotype GC/GG had increased risk (OR 4.07, $p < 0.001$)
	<i>MET</i>						rs78116323		
Song et al. 2012 [128]		CG	Korea	To investigate the genetic contribution of ALO15 to the development of AERD.	195 HC 171 AERD (49 NP) 229 ATA (9 NP)	Blood	<i>ALOX15</i>	rs34104097 rs7220870 rs2664592	The patients carrying haplotype 1 (GCG) of Rs34104097, Rs7220870, and Rs2664592 showed a significantly higher total eosinophil count compared to the other haplotypes ($p = 0.008$) in the AERD group
Szabo et al. 2013 [49]		CG	Hungary	To determine whether TNFa -308G>A SMP has a role in the genetic predisposition to CRS in a Hungarian population.	169 HC 326 CRSwNP 49 CRSsNP	Buccal cells	<i>TNF</i>	rs1800629	There was a significantly higher carriage rate of the rare A allele-containing genotypes among the AIA CRSwNP patients
Szabo et al. 2015 [48]		CG	Hungary	To examine whether the association between TNFa -308A allele and AIA CRSwNP is due to this allele or to the presence of the complete 8.1 ancestral haplogroup (AH) in chromosome 6.	169 HC 244 CRSwNP 57 CRSsNP	Buccal cells	<i>TNF</i>	rs1800629	Carriers of 8.1 AH carried all 4 studied SNPs in homozygotic or heterozygotic forms. This AH is significantly associated with CRSwNP ($p = 0.014$)
	<i>AGER</i>						rs1800625		
	<i>HSP70-2</i>						rs1061581		
	<i>LTA</i>						rs909253		
Tewfik et al. 2009 [31]		CG	USA	To investigate whether polymorphisms in the genes encoding key TLR signaling molecules might be associated with total serum IgE levels.	154 CRSwNP 27 CRSsNP	Blood	<i>TLR11</i>	rs4286521 rs4833095	Blood IgE levels have been shown to be raised in patients with CRSwNP The C allele of rs1461567, the G allele of rs4251513, and the A allele of rs4251559 of the IRAK4 gene are associated with high serum levels of IgE in the NP patients.
	rs5743611 rs5743594								
	rs4833103								
	<i>TLR2</i>						rs13150331 rs4696480		
							rs1898830 rs4696483		
							rs3804100 rs5743704		
							rs5743708 rs7656411		
							rs1339 rs17030340		
							rs2289318 rs7695605		
	<i>TLR3</i>						rs956239 rs4861699		
							rs5743305 rs7657186		
							rs6552950 rs3775296		
rs35140061 rs7668666									
rs3775292 rs35311343									
rs5743317 rs3775291									
rs5743318 rs10025405									
rs4862633 rs4608848									
rs6857595 rs1519309									
<i>TLR4</i>	rs10983754 rs10759930								
	rs10759932 rs2149356								
	rs4986790 rs4986791								
	rs11536889 rs11536898								
	rs1554973 rs7860896								
	rs7037225 rs2183016								
<i>TLR6</i>	rs5743810 rs5743808								
	rs5743794 rs5743788								
	rs6833914								

Supplementary Table 1. Selected genetic studies.

							<i>TLR9</i>	rs352162 rs352140 rs5743836 rs187084 rs352143 rs11717574	
							<i>TLR10</i>	rs4513579 rs11466657 rs11096955 rs11466652 rs10856839 rs7653908	
							<i>CD14</i>	rs7721577_TC rs4914_CG rs2569190_GA rs2569193_GA rs2563310_GA	
							<i>MD2/LY96</i>	rs1905045_TC rs16938755_TC rs11786591_CT rs6472812_GA rs10504554_TC rs17226566_TC rs12544736_TG rs16938766_GC	
							<i>MYD88</i>	rs2239621 rs4988453 rs7744 rs6767684 rs6796045	
							<i>IRAK4</i>	rs11182250 rs1461567 rs4251580 rs4251520 rs4251559 rs17121283 rs6582484 rs4251459 620-1delAC rs4251487 821delT rs4251583 T877C rs4251513 A1188+520G G1189-1T rs4251545	
							<i>TRAF6</i>	rs3740961 rs5030437 rs5030416 rs5030411 rs331455	
Tournas et al. 2010 [129]		GWAS	Canada	To verify an association between p73 and CRS.	196 HC 154 CRSwNP 52 CRSsNP	Blood	<i>P73</i>	rs3765731 rs3765692	The A allele of rs3765731 was differentially expressed in NP when compared to HC (p=0.037). The A allele has a protective effect: AA+AG vs GG OR 0.5391, p=0.0036.
Wang et al. 2000 [68]		CG	USA	To determine whether mutations in the CFTR gene, which is responsible for CF, predispose to CRS.	123 HC 147 CRS	Blood	<i>CFTR</i>	ΔF508 G542X N1303K	Only 11 patients had one of the selected mutations in the CFTR gene.
Wang et al. 2008 [130]		CG	Taiwan	To investigate the role of MMP2 tagging SNPs and promoter functional polymorphism in the	136 HC 136 CRSwNP	Blood	<i>MMP2</i>	rs2438656 rs857403 rs1030868 rs1477017	rs857403 T allele was associated with increased risk (OR 2.07 p=0.03) but it could not be replicated with additional controls.

Supplementary Table 1. Selected genetic studies.

				development of NP.				rs1053605 rs9302671 rs2241145 rs2241146 rs243849 rs12599775 rs243847 rs243844 rs243840 rs2287076 rs11639960 rs243832 rs7201	
Wang et al. 2010 [131]		CG	Taiwan	To investigate the role of MMP9 tagging SNPs and promoter functional polymorphism in the development of NP.	730 HC 203 CRSwNP	Blood	MMP9	rs3918242 rs2664538 rs3787268 rs2274756	The T allele of promoter SNP rs3918242 was associated with CRSwNP under the dominant (nominal p = 0.023, empirical p = 0.022, OR = 1.62) and additive models (nominal p = 0.012, empirical p = 0.011, OR = 1.60). The A allele of rs2274756 has a nominal p value of 0.034 under the dominant model and 0.020 under the additive model. The most significant haplotype was TGGA p=0.0045
Wang et al. 2013 [132]		CG	Taiwan	To investigate the relative expression of MMPs in the non-recurrent and recurrent NP as compared to control individuals.	31 HC 30 CRSwNP	Blood	MMP2 MMP9	rs243865 rs3918242	Genetic polymorphisms of MMP-2 and MMP-9 functional promoters were not associated with the recurrence of NP.
Yazdani et al. 2012 [133]		CG	Iran	To investigate the association between the polymorphism C-159T in CD14 gene and NP.	87 HC 107 CRSwNP	Blood	CD14	rs946564423	Significant association of the C allele in NP patients (or 1.88, p=0.04)
Yea et al. 2006 [45]		CG	Korea	To investigate the relationship between an IL-4 promoter polymorphism and NP.	70 HC 106 CRS 61 CRSwNP	Blood	IL4	-590C/T	The presence of T allele was associated with reduced risk of NP (OR TT 0.529, p=0.028)
Zhai et al. 2007 [26]		CG	China	To explore a potential association between NP and polymorphisms at loci for HLA-DR and HLA-DQ.	81 HC 30 NP	Blood	HLA-DR HLA-DQ	*04 *07 *08 *09 *10 *11 *12 *13 *14 *15 *16 *02 *04 *05 *06 *07 *08 *09	Frequency of allele was significantly higher in patients for DR*09 and -*16 and DQ-*08 and -*09. DQ*07 frequency was lower in patients.
Zhang et al. 2008 [134]		CG	China	To examine whether there is an association between Clara cell 10kDa protein (CC10)+38A>G, plasma CC10 levels and CRS.	180 HC 90 CRSwNP 130 CRSsNP	Blood	CC10 (SCGB1A1)	+38A>G	No association
Zhang et al. 2011 [135]		CG	Canada	To determine whether polymorphisms in gene regulating NO synthesis	187 HC 154 CRSwNP	Blood	NOS1	rs1004356 rs1483757	Two SNPs in the NOS1 gene (rs1483757, p=0.0023, OR 0.62; rs9658281, p =0.0129, O 0.66) remained

Supplementary Table 1. Selected genetic studies.

				are associated with CRS.	52 CRSsNP			rs545654 rs9658281	significant after correction for multiple testing. Homozygote allele C (p=0.0017; OR 0.28) in rs1483757 locus increased the risk.
							<i>NOS1AP</i>	rs10458392 rs10919117 rs12022557 rs12047527 rs12061249 rs3923367 rs4657164 rs6676638 rs6677052 rs6677606 rs7416392 rs7546353 rs6681981 rs8179404	rs12047527 in NOS1AP showed significant association (p<0.05) with CRS
Zhang et al. 2012a [40]		CG	China	To replicate and extend genetic association studies in CRS in a Chinese population.	315 HC 306 CRSwNP 332 CRSsNP	Blood	<i>PARS2</i>	rs2873551	Rs4532099 SNP in RYBP increased the risk of CRSwNP (OR 2.76, p=3.2x10 ⁻⁶).
							<i>IL22RA1</i>	rs4292900 rs4648936 rs16829225	Selected SNPs in AOA and IRAK4 were associated with a reduced risk of CRS (OR 0.60-0.79, p<0.05)
							<i>TNFRSF1B</i>	rs235214 rs496888 rs652625 rs7550488	
							<i>TRIP12</i>	rs1035833	
							<i>IL1RL1</i>	rs13431828 rs10204137	
							<i>IL1A</i>	rs17561 rs2856838 rs2048874 rs1800587	
							<i>FAM79B</i>	rs13059863	
							<i>RYBP</i>	rs4532099	
							<i>TSLP</i>	rs3806932 rs2289276	
							<i>LAMA2</i>	rs2571584	
							<i>TNFAIP3</i>	rs3757173 rs5029938	
							<i>LAMB1</i>	rs4727695	
							<i>AOAH</i>	rs4504543	
							<i>MET</i>	rs38850	
							<i>RAC1</i>	rs836479	
							<i>CACNA2D1</i>	rs6972720	
							<i>KIAA1456</i>	rs11779957	

Supplementary Table 1. Selected genetic studies.

							<i>MSRA</i>	rs7001821	
							<i>MUSK</i>	rs10817091	
							<i>PDGFD</i>	rs12574463	
							<i>NOS1</i>	rs1483757	
							<i>NAV3</i>	rs1726427	
							<i>IRAK4</i>	rs4251431 rs6582484 rs1461567 rs3794262	
							<i>SERPINA1</i>	rs1243168 rs4900229	
							<i>UBE3A</i>	rs1557871	
							<i>SLC13A3</i>	rs393990	
							<i>CACNA1I</i>	rs3788568	
Zhang et al. 2013a [136]		CG	China	To examine association between specific SNPs in/around the FOXP3 and EB13 genes and susceptibility to CRS	315 HC 306 CRSwNP 332 CRSsNP	Blood	<i>EB13</i>	rs428253 rs6613 rs353698 rs2302164	Risk analysis showed rs428253 of EB13 gene to play a protective role among both CRSsNP (GG/CC) and CRSwNP (CG/CC) subjects. Haplotype analysis of the FOXP3 gene region further indicated that CRS risk was higher in individuals carrying the haplotype GG in rs2294018–rs2232365 block, compared with wild-type AG haplotype
							<i>FOXP3</i>	rs2294018 rs3060515 rs2232365 rs3761548 rs4824747	
Zhang et al. 2013b [137]		CG	China	To explore associations between SNPs in/around the TSLP gene and development of CRS	315 HC 306 CRSwNP 332 CRSsNP	Blood	<i>TSLP</i>	rs1545169 rs764917 rs12653736 rs1837253 rs12654933 rs10455025 rs11466741 rs13156086 rs6886755 rs252706 rs2416259	SNPs rs252706 (AA genotype: P=0.012, OR 0.552) and rs764917 (CA genotype: P=0.001, OR 0.182) displayed protective roles among CRSwNP, but not CRSsNP, subjects.
Zielinska et al. 2012 [138]		CG	Poland	To investigate the association between LF and OSF2 polymorphisms with the risk of CRSwNP in Poland	200 HC 195 CRSwNP	Blood	<i>LTF</i>	rs1126478	Rs1126478 LF (OR 4.78; 95% CI 3.07–7.24), the -33C/G OSF2 (OR 3.48; 95% CI 2.19–5.52) and the rs3829365 OSF2 (OR 16.45; 95% CI 6.71–40.30) genotypes were associated with an increased risk of CRSwNP.
							<i>fgOSF2</i>	rs3829365 -33C/G	

Supplementary Table 2. Selected epigenetic studies.

Reference	Objective	Tissue	Epigenetic assay	Population	Significant findings
Callejas-Diaz et al. 2020 [84]	To identify which key mRNA and miRs are regulating in vitro mucociliary differentiation of human adult basal stem cells under pathological and healthy conditions.	NP, inferior turbinate mucosa (ITM; control)	miRNA	Spain	Transcriptome related to ciliogenesis and cilia function is significantly impaired during differentiation of CRSwNP epithelium due to an altered expression of microRNAs, particularly of those belonging to mir-34 and mi-449 families
Cheong et al. 2011 [76]	To analyze the genome-wide DNA methylation levels in nasal polyps from patients with AIA.	NP, buffy coats	Genome-wide DNA methylation	China	332 loci in 296 genes were hypermethylated in AIA vs ATA. These genes are involved in ectoderm development, hemostasis, and wound healing. 158 loci in 141 genes were hypomethylated in AIA vs ATA. Relevant pathways were lymphocyte proliferation, cell proliferation, leukocyte activation, and immune response.
Cho et al. 2012 [75]	To study the effect of trichostatin A (TSA), a histone deacetylase inhibitor, on TGFβ1-induced myofibroblast differentiation and ECM accumulation in NP fibroblasts.	NP, ITM	Histone acetylation control	Korea	The expression levels of HDAC2, α-SMA and TGF-β1 were increased in NP compared to normal tissues. TSA induced hyperacetylation of histones, inhibiting them. HDAC inhibition is associated with myofibroblast differentiation and ECM accumulation in NP.
Cho et al. 2013 [75]	To investigate the inhibitory effect of TSA on myofibroblast differentiation and ECM production in nasal polyp organ cultures.	NP tissue cultures	Histone acetylation control	Korea	TSA inhibited HDAC and induced hyperacetylation of histones H4
Kidoguchi et al. 2018 [77]	To investigate the methylation levels at 3 CpG sites in the proximal PLAT promoter and their effects on gene expression in NP tissue.	NP, ITM	DNA methylation	Japan	Methylation of -618, -121, and -105 CpGs was significantly higher in NP. <i>PLAT</i> expression was lower ($p > 0.001$). The methylation changes at -618 site showed a negative correlation with the gene expression changes between NP and ITM ($r = -.65$, $p < 0.01$).
Kim et al. 2018 [78]	To elucidate whether DNA methylation of specific genes is involved in the development of NP.	NP, ITM	DNA methylation	Korea	The promoter regions of 10 and 30 genes were hypermethylated and hypomethylated, respectively, in NP samples compared with controls. The top four genes with altered hypomethylation in NP tissues were <i>KRT19</i> , <i>NR2F2</i> , <i>ADAMTS1</i> and <i>ZNF222</i> .
Kim et al. 2019 [79]	To investigate the expression and distribution of FZD5 and the role of eosinophil infiltration in CRSwNP pathogenesis.	NP, uncinated process tissue	Methylation profiling	Korea	397 and 387 genes were differentially hypermethylated and 399 and 208 genes were hypomethylated in the E-CRSwNP and NE-CRSwNP groups, respectively, compared to the control tissues. Most of the differentially methylated genes were associated with cancer pathways. FZD5 was significantly hypomethylated in the E-CRSwNP compared to the NE-CRSwNP group.
Li et al. 2019a [80]	To determine whether there was any association between abnormal DNA methylation of TSLP gene and CRS	NP, ethmoid mucosa (CRSsNP) patients	DNA methylation	China	There was an increase in methylation ratios of 4 CpGs (2, 22, 23, 24) of TSLP gene had increased in the CRSwNP patients compared to the CRSsNP and

Supplementary Table 2. Selected epigenetic studies.

	pathogenesis.	and ITM			control subjects, significantly related to disease status (p<0.02)
Li et al. 2019b [81]	To determine whether there was any association between abnormal DNA methylation of IL8 promoter and CRS pathogenesis.	NP, ethmoid mucosa (CRSSNP) patients and ITM	DNA methylation	China	Three CpGs (-116, -106, -31) were significantly hypomethylated in the CRSSNP group compared with CRSSNP and HC.
Liu et al. 2018 [85]	To study the role of miR124 in CRSwNP.	NP, ITM	miRNA	China	MiR124 expression was reduced in NP tissues, which negatively correlated with the expression of AHR. This may be critical to the development of inflammatory response in CRSwNP.
Liu et al. 2019 [83]	To characterize the transcriptome profiles of mRNAs and lncRNAs in patients with CRSwNP.	GEO datasets, blood samples	lncRNA	China	A total of 265 differentially expressed lncRNAs were obtained, including 56 upregulated and 209 downregulated genes.
Luo et al. 2017 [86]	To test whether miR-17-92 cluster is associated with suppressing IL-10 in peripheral DC.	Blood samples	miRNA	China	A negative correlation was found between expression of IL-10 and miR-19a in DC from NP patients. miR19-1 was upregulated while miR-17, -18a, -19b, -20a and -92a showed no differences between NP and HC.
Ma et al. 2015 [88]	To investigate miRNAs expression profiles of peripheral blood DCs in CRS patients.	Blood samples	miRNA	China	There were 31 miRNAs changed in all CRS patients with respect to HC, and 49 miRNA that changed exclusively in CRSwNP. miR-210-3p, miR-125b-5p, and miR-150-5p were upregulated in CRS, while miR-708-5p and miR-126-3p were downregulated.
Ma et al. 2018 [87]	To investigate the effects and mechanism of miR-150-5p to promote the development of CRS via the DC-Th axis.	Blood samples	miRNA	China	miR-150-5p was upregulated in DCs from CRS patients compared with HC, and DCs Promote Naïve T Cells Proliferation. MiR-150-5p further regulated EGR2 and inhibited DCs, leading to an abnormal DC-Th axis.
Qing et al. 2019 [89]	To investigate the mechanisms between the miR-142-3p and TNF- α activation in vitro and in vivo	NP, ITM	miRNA	China	miR-142-3p may participate in the regulation of the body's inflammatory response through the LPS-TLR-TNF- α signaling pathway in CRSwNP.
Seiberling et al. 2012 [95]	To determine the presence of 5-bromo-cytosine, 5-chloro-cytosine and methylated cytosine in CRSwNP.	NP, posterior ethmoid tissue (HC)	DNA modification	USA	The levels of 5-Bromocytosine were significantly higher in polyps (p=0.007). Aberrant methylation patterns in polyp eosinophils may help explain the pathogenesis of CRSwNP.
Tian et al. 2012 [96]	To explore the profiling of tandem alternative polyA (APA) sites in NP.	NP, uncinated process mucosa	Genome-wide polyadenylation site sequencing	China	There was a switching of 3'UTR lengths in NP compared with nasal uncinated process mucosa from the same patient. 105 genes were switched to distal polyA sites in the nasal polyps and 90 genes were switched to proximal poly(A) sites. Besides, 213 genes were upregulated in NP while 414 genes were downregulated.
Xuan et al. 2019 [90]	To evaluate miRNAs profiles and relevant biological pathways in CRSwNP and	Nasal mucosa	miRNA array	China	24 miRNAs showed differential expression. 5 miRNAs (miR-210-5p, miR-3178, miR-585-3p, miR-3146, and

Supplementary Table 2. Selected epigenetic studies.

	control subjects.				miR-320e) were significantly upregulated ($p < 0.05$, fold change >2), and 19 miRNAs, including miR-32-3p, miR-1299, miR-3196, miR-3924, miR-548e-3p, miR-3184-5p, miR-375, miR-23a-5p, miR-377-5p, miR-574-5p, miR-3149, miR-500a-5p, miR-125b-2-3p, miR-1914-5p, miR-532-3p, miR-612, miR-1298-5p, miR-1226-3p, and miR-668-3p, were significantly downregulated in CRSwNP tissue ($p < 0.05$, fold change <0.5).
Yan et al. 2020 [91]	To examine human neutrophil elastase-induced MUC5AC overexpression in CRS via miR-146a.	NP, uncinated process mucosa	miRNA	China	EGFR is a target of miR-146a. This miRNA is downregulated in NP reducing the inhibition of EGFR, and therefore MUC5AC expression levels were increased.
Yu et al. 2018 [92]	To evaluate the roles of TGF β 1 and miR-663 in the pathogenesis of NP in children.	Nasal mucosa, peripheral blood eosinophils (PBE)	miRNA	China	The expression of miR-633 was significantly reduced in polyps and PBE from CRS patients, while <i>TGFB1</i> mRNA was significantly increased. miR-633 binds to the 3' UTR of <i>TGFB1</i> and regulated its expression.
Zhang et al. 2012b [94]	To determine the pattern of expression and biological role of miRNAs in CRS.	NP, ethmoidal mucosa, inferior turbinate tissue	miRNA	China	miR-125b was upregulated in CRSwNP when compared to CRSsNP. This may enhance type I IFN expression through suppressing 4E-BP1 protein expression in airway epithelial cells.
Zhang et al. 2012c [97]	To investigate the expression of miRNA machinery components in CRS.	NP, ethmoid sinus mucosa	mRNA expression	China	PACT mRNA expression was found to be upregulated in CRSwNP as compared with controls. The rest of the miRNA machinery components including Drosha, Dicer, TRBP, FXR1 and E1F2C2, showed no differences between patients and controls.
Zheng et al. 2015 [82]	To identify whether DNA methylation plays a role in the pathogenesis of NP.	NP, ITM	DNA methylation	China	198 genes had a differential methylated signal in their promoter region when comparing NP samples with ITM samples. The four most changed genes were <i>COL18A1</i> , <i>EP300</i> , <i>GNAS</i> and <i>SMURF1</i> .
Zhou et al. 2020 [93]	To explore the pathogenesis of CRSwNP from the perspective of genes.	CRSwNP datasets. NP, nasal mucosa (HC)	Functional enrichment analysis, including non-coding RNAs	China	Two clusters of genes, lncRNAs and miRNAs were found to be related to CRSwNP. Main miRNA involves were: miR-130a, miR-27a-3p, miR-193-3p, miR-29a-3p, miR-18b-5p, miR-138-5p, and miR-25-3p.

Supplementary Table 3.

Functional Category	Enrichment FDR	Genes in list	Total genes	Genes
Cytokine-mediated signaling pathway	2.63e-16	29	950	IL1B IL1RN IL22RA1 CCL11 IRAK4 TSLP EBI3 IL1RL1 FCER1G IL1A PPARG TNF NOS2 ALOX5 MMP2 MMP9 IL10 IL33 ALOX15 CIITA HLA-DRB3 HLA-DRB1 HLA- DQA1 HLA-DRB5 HLA-DRA HLA-C HLA-B HLA-A HLA-DRB4
Defense response	1.10e-15	38	2062	NOS2 IL33 FCER1G PTGDR CD14 CCL11 CIITA LTF IL1B IL10 TNF HLA-A ALOX5 FOXP3 IL1A IL1RL1 PPARG ALOX5AP AOAH IL1RN IL22RA1 MS4A2 ADORA1 CYSLTR1 SERPINA1 IRAK4 AGER TSLP HLA- DRB1 MMP9 ALOX15 HLA- DRB3 HLA-DQA1 HLA-DRB5 HLA- DRA HLA-C HLA-B HLA-DRB4
Inflammatory response	1.10e-15	27	856	IL33 PTGDR CD14 CCL11 CIITA NOS2 IL1B IL10 TNF ALOX5 FOXP3 IL1A IL1RL1 PPARG ALOX5AP AOAH IL1RN MS4A2 FCER1G ADORA1 CYSLTR1 SERPINA1 AGER TSLP HLA-DRB1 MMP9 ALOX15
Response to stress	7.689e-15	52	4507	NOS2 MMP2 CAT IL1B HSPA2 IL1RN IL33 TRIP12 FANCC FCER1G PTGDR CD14 CCL11 MSRA CIITA CFTR LTF DCBLD2 TP73 NOS1 MMP9 IL1A IL10 TNF HLA-A IFRD1 ALOX5 FOXP3 IL1RL1 PPARG ALOX5AP AOAH IL22RA1 MS4A2 ALOX15 ADORA1 ADRB2 CYSLTR1 SERPINA1 IRAK4 MT-CO2 AGER MET TSLP HLA- DRB1 HLA-DRB3 HLA-DQA1 HLA- DRB5 HLA-DRA HLA-C HLA-B HLA-DRB4

Supplementary Table 3.

Response to cytokine	1.04e-13	30	1372	IL1B IL1RN IL22RA1 CCL11 IRAK4 TSLP CIITA NOS2 EBI3 IL1RL1 FCER1G CD14 IL1A PPARG ALOX15 TNF ALOX5 MMP2 MMP9 IL10 IL33 HLA-DRB3 HLA- DRB1 HLA-DQA1 HLA-DRB5 HLA- DRA HLA-C HLA-B HLA-A HLA-DRB4
Immune system process	1.124e-13	45	3539	RUNX2 IL1B IL1RN CD8A FCER1G CD14 CCL11 ACE LTF FOXP3 PPARG IL10 IL33 CIITA HLA-DRB1 HLA- DRA AGER TNF HLA-B HLA-A NOS2 MMP9 EBI3 TAPBP IL1A IL1RL1 MS4A2 FANCC ALOX15 ADORA1 CYSLTR1 LTA HLA-DQB1 HLA- DRB3 HLA-DQA1 IRAK4 HLA-DRB5 HLA-C HLA-DRB4 TSLP ALOX5 TP73 CAT FCER1A SERPINA1
Cellular response to cytokine stimulus	1.121e-13	29	1278	IL1B IL1RN IL22RA1 CCL11 IRAK4 TSLP NOS2 EBI3 IL1RL1 FCER1G CIITA IL1A PPARG ALOX15 TNF ALOX5 MMP2 MMP9 IL10 IL33 HLA-DRB3 HLA- DRB1 HLA-DQA1 HLA-DRB5 HLA- DRA HLA-C HLA-B HLA-A HLA-DRB4
Immune response	1.41e-13	39	2602	CD8A FCER1G CD14 CCL11 IL1B LTF FOXP3 IL10 CIITA AGER TNF HLA-B HLA-A NOS2 IL1A IL1RL1 IL1RN IL33 MS4A2 ALOX15 CYSLTR1 LTA HLA-DQB1 HLA-DRB3 HLA- DRB1 HLA-DQA1 IRAK4 HLA-DRB5 HLA-DRA HLA-C HLA-DRB4 PPARG ALOX5 MMP9 EBI3 TAPBP CAT FCER1A SERPINA1
Cell surface receptor signaling pathway	2.29e-13	43	3287	MUSK MET IL1B IL1RN IL22RA1 CD8A FCER1G CD14 CCL11 ANXA4 IRAK4 LTF

Supplementary Table 3.

				TSLP TNF FOXP3 EBI3 IL1A IL1RL1 RUNX2 IL33 MS4A2 ADORA1 ADRB2 CYSLTR1 MMP9 PPARG NOS2 ALOX5 MMP2 IL10 ALOX15 CIITA FCER1A HLA- DRB3 HLA-DRB1 HLA-DQA1 HLA- DRB5 HLA-DRA AGER HLA-C HLA-B HLA-A HLA-DRB4
Cellular response to chemical stimulus	4.85e-13	44	3536	IL1B HSPA2 PPARG IL1RN IL22RA1 FANCC FCER1G CD14 CCL11 MSRA IRAK4 CFTR LTF MMP9 ALOX5AP TSLP ALOX15 AGER TNF NOS2 MMP2 NOS1 EBI3 IL1RL1 CAT RUNX2 IL10 PTGDR ADRB2 CIITA LTC4S MET IL1A HLA-DRB1 ALOX5 IL33 HLA- DRB3 HLA-DQA1 HLA-DRB5 HLA- DRA HLA-C HLA-B HLA-A HLA-DRB4
Cellular response to organic substance	5.90e-12	39	2938	IL1B HSPA2 PPARG IL1RN IL22RA1 CD14 CCL11 IRAK4 CFTR LTF TSLP AGER TNF NOS2 MMP2 NOS1 EBI3 IL1RL1 CAT RUNX2 IL10 FCER1G PTGDR ADRB2 CIITA IL1A ALOX15 HLA- DRB1 ALOX5 MMP9 IL33 HLA- DRB3 HLA-DQA1 HLA-DRB5 HLA- DRA HLA-C HLA-B HLA-A HLA-DRB4
Response to organic substance	1.593e-11	42	3547	NOS2 NOS1 IL1B HSPA2 PPARG IL1RN IL22RA1 CD14 CCL11 IRAK4 CFTR LTF IL10 TSLP HLCS CIITA AGER TNF TBXAS1 TP73 MMP2 MMP9 EBI3 IL1RL1 CAT RUNX2 FCER1G PTGDR ADRB2 IL1A ALOX15 HLA-DRB1 ALOX5 IL33 HLA-DRB3 HLA-DQA1 HLA- DRB5 HLA-DRA HLA-C HLA-B HLA-A HLA-DRB4
Regulation of immune system process	2.414e-10	30	1909	FCER1G CD14 IL1B ACE FOXP3

Supplementary Table 3.

				IL10 IL33 AGER TNF HLA-B HLA-A IL1RL1 MS4A2 ALOX15 ADORA1 LTF PPARG TSLP HLA-DRB1 TP73 EBI3 CD8A FCER1A HLA- DRB3 HLA-DQA1 IRAK4 HLA-DRB5 HLA-DRA HLA-C HLA-DRB4
Regulation of immune response	5.63e-10	25	1325	FCER1G CD14 IL1B FOXP3 IL10 AGER TNF HLA-B HLA-A IL1RL1 IL33 MS4A2 ALOX15 LTF PPARG CD8A FCER1A HLA- DRB3 HLA-DRB1 HLA-DQA1 IRAK4 HLA-DRB5 HLA- DRA HLA-C HLA- DRB4
Regulation of response to stimulus	1.05e-09	46	4820	IL1B IL1RN IL33 FCER1G CD14 CCL11 IRAK4 NOS2 LTF FOXP3 MET IL10 TSLP ADRB2 AGER TNF HLA-B HLA-A CFTR TP73 NOS1 EBI3 IL1A IL1RL1 CAT RUNX2 PPARG ALOX5AP AOAH MS4A2 ALOX15 ADORA1 LTA RGS7BP MMP9 TRIP12 HLA-DRB1 NOS1AP CD8A FCER1A HLA- DRB3 HLA-DQA1 HLA-DRB5 HLA- DRA HLA-C HLA- DRB4
Cytokine secretion	3.78e-09	13	285	CD14 NOS2 FOXP3 IL1A IL10 IL33 TNF IL1RL1 IL1B AGER TSLP ANXA4 HLA-DRB1
Cellular response to interferon-gamma	4.23e-09	13	289	CCL11 NOS2 CIITA PPARG HLA-DRB3 HLA- DRB1 HLA-DQA1 HLA-DRB5 HLA- DRA HLA-C HLA-B HLA-A HLA-DRB4
Interferon-gamma-mediated signaling pathway	4.76e-09	11	178	PPARG CIITA HLA-DRB3 HLA- DRB1 HLA-DQA1 HLA-DRB5 HLA- DRA HLA-C HLA-B HLA-A HLA-DRB4
Regulation of inflammatory response	6.81e-09	15	448	IL33 NOS2 FOXP3 IL1RL1 IL1B PPARG ALOX5AP AOA IL10 FCER1G ADORA1 AGER TSLP TNF MMP9
Antigen processing and presentation	9.27e-09	14	384	FCER1G HLA- DRB1 HLA-DRA HLA-B TAPBP

Supplementary Table 3.

				HLA-DQB1 HLA-DRB3 HLA-DQA1 HLA-DRB5 HLA-C HLA-A HLA-DRB4 CD8A ACE
Response to interferon-gamma	9.27e-09	13	312	CCL11 CIITA NOS2 PPARG HLA-DRB3 HLA-DRB1 HLA-DQA1 HLA-DRB5 HLA-DRA HLA-C HLA-B HLA-A HLA-DRB4
Regulation of cytokine secretion	1.269e-08	12	257	CD14 FOXP3 IL1A IL10 IL33 TNF IL1RL1 IL1B AGER TSLP ANXA4 HLA-DRB1
Positive regulation of response to stimulus	1.51e-08	32	2621	IL1B IL1RN IL33 FCER1G CD14 CCL11 IRAK4 LTF IL10 TSLP ADRB2 TNF HLA-B CFTR FOXP3 TP73 NOS1 IL1RL1 CAT ALOX5AP ADORA1 AGER MMP9 MET ALOX15 HLA-DRB1 NOS1AP HLA-DRB3 HLA-DQA1 HLA-DRB5 HLA-DRA HLA-DRB4
Cytokine production	4.97e-08	19	925	IL1B IL1RN CD14 NOS2 LTF FOXP3 IL1A IL10 IL33 TSLP AGER TNF IL1RL1 FCER1G IRAK4 NAV3 ANXA4 HLA-DRB1 EBI3
Interleukin-6 production	4.97e-08	10	172	IL1B IL1RN NOS2 IL10 TNF FOXP3 IL33 FCER1G AGER TSLP
Secretion	7.35e-08	26	1861	CACNA1I CD14 NOS2 FOXP3 IL1A IL1B IL10 IL33 ACE TNF CFTR IL1RL1 IL1RN FCER1G ADORA1 MT-CO2 AGER TSLP ANXA4 HLA-DRB1 LTF ALOX5 MMP9 CAT SERPINA1 HLA-C
Positive regulation of transport	7.35e-08	20	1069	CACNA1I FCER1G CD14 CFTR IL1A IL10 IL33 TNF IL1RL1 IL1B HSPA2 PPARG ADORA1 AGER ADRB2 TSLP NOS1 HLA-DRB1 NOS1AP TP73
Regulation of cytokine production	8.36e-08	18	852	IL1B IL1RN CD14 NOS2 LTF FOXP3 IL1A IL10 IL33 TSLP AGER TNF IL1RL1 FCER1G NAV3 ANXA4 HLA-DRB1 EBI3
Regulation of defense response	8.789e-08	19	968	IL33 CD14 NOS2 HLA-A FOXP3

Supplementary Table 3.

				IL1RL1 IL1B PPARG ALOX5AP AOAH IL10 FCER1G ADORA1 AGER LTF TSLP TNF MMP9 IRAK4
Regulation of peptide secretion	9.132e-08	15	559	CD14 FOXP3 IL1A IL1B IL10 IL33 TNF CFTR IL1RL1 ADORA1 AGER NOS2 TSLP ANXA4 HLA-DRB1