

Table SI. Complete list of genotyped variants

Gene name (symbol)	Protein function	Specific relevance in AD	Position	Variant (consequence)	MAF % AD subjects	MAF % controls	p-value (OR; 95% CI)
Claudin 1 ( <i>CLDN1</i> )	Transmembrane protein in tight junction strand.	Expression reduced in AD. The reduction seems to diminish the integrity of tight junctions and correlates inversely with Th2 cytokines. Certain single-nucleotide polymorphisms (SNPs) of <i>CLDN1</i> are associated with increased risk of AD in North American populations. Defects may be involved in the higher susceptibility for Herpes simplex virus (HSV) infections seen in AD patients. Deficiency increases transepidermal water loss (TEWL) in mice, leading to a death of the <i>CLDN1</i> -deficient mice in 1 day. Clobetasol reduces <i>CLDN1</i> expression (21, 22, S1–S3).	190046808	rs16865373	3.483	4.304	0.286 (0.800; 0.531–1.205)
			190034709	rs3732923	38.430	42.550	0.032 (0.843; 0.721–0.986)
			190026860	rs3774032	16.850	17.200	0.812 (0.976; 0.796–1.196)
			190026083	rs141397566	0.112	0.040	0.467 (2.797; 0.175–44.82)
			190030679	rs140846629	2.472	2.454	0.976 (1.008; 0.615–1.651)
Claudin 4 ( <i>CLDN4</i> )	Transmembrane protein in tight junction strand.	Clobetasol and histamine seem to reduce <i>CLDN4</i> expression (S3, S4)	73245485	rs72466475	0.449	0.443	0.980 (1.015; 0.322–3.204)
Claudin 20 ( <i>CLDN20</i> )	Transmembrane protein in tight junction strand.		155597134	rs34434986	6.629	6.034	0.52 (1.11; 0.808–1.524)
Claudin 23 ( <i>CLDN23</i> )	Transmembrane protein in tight junction strand.	Expression seems to be reduced in AD (21).	8560107	rs61755871	0.246	0.041	0.143 (6.03; 0.545–66.67)
			8560536	rs12548737	13.820	15.570	0.215 (0.869; 0.695–1.086)
Filaggrin ( <i>FLG</i> )			152285076	rs558269137 ( <i>2282del4</i> )	3.378	1.327	0.00016 (2.657; 1.601–4.411)
			152284303	rs200360684 ( <i>S1020X</i> )	0.225	0.080	0.304 (2.801; 0.393–19.95)
			152285861	rs61816761 ( <i>R501X</i> )	0.899	0.081	0.0022 (11.29; 2.387–53.35)
			152277622	rs150597413 ( <i>S3247X</i> )	0	0.040	–
			152274016	rs12730241 ( <i>12-repeat allele</i> )	20.110	17.900	0.148 (1.152; 0.951–1.397)
			152282102	NA	0	0.040	–
			152280023	rs138726443 ( <i>R2447X</i> )	1.348	0.443	0.007239 (3.099; 1.357–7.074)
			152285555	rs137995883 ( <i>V603M</i> )	0	0.041	–
Filaggrin 2 ( <i>FLG2</i> )	Exact function of the protein not known.	Expression seems to follow that of filaggrin. <i>FLG2</i> mutations are associated with more persistent AD in African-American patients (S5, S6).	152329370	rs145678751	0.225	0.081	0.305 (2.797; 0.393–19.91)
			152329758	rs61749580	0	0.040	–
			152323132	rs12568784	14.040	14.480	0.758 (0.967; 0.78–1.199)
			152329852	rs6587667	0.903	0.617	0.381 (1.472; 0.62–3.497)
			152326321	NA	0	0.040	–
			152329369	rs2282302	20.720	20.260	0.765 (1.03; 0.849–1.249)
Involucrin ( <i>IVL</i> )	A component of the crosslinked envelope in terminally differentiated keratinocytes. Crosslinked to membrane proteins, binds to loricrin.	Topical betamethasone therapy decreases <i>IVL</i> expression compared with calcineurin inhibitors, a possible explanation for the lack of functional restoration of stratum corneum layers seen after treatment with betamethasone. UV therapy seems to normalize <i>IVL</i> expression instead (S7, S8).	152883573	rs139703221	0	0.040	–
Junctional adhesion molecule A; F11 receptor ( <i>JAM1</i> ; <i>F11R</i> )	An important regulator of tight junction assembly.		160970508	rs116727809	0.112	0.201	0.594 (0.558; 0.065–4.786)
Loricrin ( <i>LOR</i> )	An important component of the cornified cell envelope found in terminally differentiated keratinocytes.	Expression is reduced in the skin of flaky tail mouse, a mouse model of AD. Topical pimecrolimus and topical betamethasone therapies seem to normalize the expression <i>LOR</i> (S9, S10).	–	–	–	–	–
Occludin ( <i>OCLN</i> )	Transmembrane protein required for regulation of the tight junction permeability barrier.	Interacts with claudins and scaffold proteins. Expression reduced by clobetasol and histamine, and to a lesser extent also by tacrolimus. Reduced expression in the mouse model of AD, the flaky tail mouse (S3, S4, S9, S11).	68804987	rs147125035	2.247	2.496	0.679 (0.898; 0.539–1.495)
			68809916	rs373344533	0	0.040	–

Table SI. Contd.

Tight junction protein 1 (TJP1; ZO-1)	Tight junction protein that may be involved in signal transduction at cell-cell junctions.	Interacts with claudins and scaffold proteins. Histamine seems to reduce expression (S4, S11).	30008977	rs2291166	5.631	3.982	0.042 (1.434; 1.013-2.03)
			30011099	rs35392254	0.112	0.040	0.467 (2.797; 0.175-44.82)
			30010841	rs201943761	0	0.081	-
			30018627	rs2229515	9.101	9.702	0.605 (0.933; 0.719-1.212)
			30026582	rs2229517	0.225	0.886	0.061 (0.25; 0.059-1.068)

AD: atopic dermatitis; SD: standard deviation; MAF: minor allele frequency; OR: odds ratio; 95% CI: 95% confidence interval; UV: ultraviolet.

## SUPPLEMENTARY REFERENCES

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