

Electronic version:

Electronic version, part 1:

Each sample was drawn into pyrogen-free vials, and, within 2 hours, the serum was separated by centrifugation and stored at -80°C . Cytokine concentrations were determined by multiplex immunoassay using a 10-plex human cytokine kit from BioRad Laboratories (Hercules, CA, USA).(1+2) The assay was performed according to the manufacturer's instructions, and was run and analysed on a Bio-Plex 100 Suspension Array System (BioRad).

1: de Jager W, te Velthuis H, Prakken BJ, *et al.* Simultaneous detection of 15 human cytokines in a single sample of stimulated peripheral blood mononuclear cells. *Clin Diagn Lab Immunol* 2003; 10(1):133–139.

2: de Jager W, Prakken BJ, Bijlsma JW, *et al.* Improved multiplex immunoassay performance in human plasma and synovial fluid following removal of interfering heterophilic antibodies. *J Immunol Methods* 2005; 300(1–2):124–135.

Electronic version, part 2:

Genotyping was performed with TaqMan genotyping technology using the manufacturers protocol. Sequences of primers and probes are given below. Negative controls consisted of ddH₂O added to the reaction mix. And were added to the PCR plate as well.

TABLE primers and probes

SNP	Primers	Probes
IL-6 -174 G/C	GACGACCTAAGCTGCACTTTTC	CTTTAGCAT <u>C</u> GCAAGAC
(RS1800795)	GGGCTGATTGGAAACCTTATTAAGATTG	CTTTAGCAT <u>G</u> GCAAGAC

SNP	Primers	Probes
IL-6 -572 G/C (RS1800796)	GCCTTGAAGTAACTGCACGAAATT CCAGTCATCTGAGTTCTTCTGTGTT	AACAGCC <u>G</u> CTCACAG TACAACAGCCC <u>C</u> TCACAG
IL-8 -251 T/A (RS4073)	GTTCTAACACCTGCCACTCTAGTAC CATTTAAAATACTGAAGCTCCACAATTTG GT	AAGCATAACA <u>A</u> TTGATAATT AGCATACA <u>I</u> TTGATAATT
IL-10 -592 C/A (RS1800872)	CCAAGACAACACTACTAAGGCTTCT GCTGGATAGGAGGTCCCTTACTTT	CCTACTTCCCC <u>C</u> TCCCAA CCTACTTCCCC <u>T</u> TCCCAA
IL-10 -1082 G/A (RS1800896)	GCCCTTCCATTTTACTTTCCAGAGA GGTAAAGGAGCCTGGAACACATC	CCCGCCTGT <u>C</u> CTGTAG CCGCCTGT <u>A</u> CTGTAG
IL-10 +3367 G/A (RS3024495)	GCAGAGTTTGATGAAAAGACATTAGAGG AA TTGGTGGGAGAACACAGACATTTAA	CTCTCACCG <u>T</u> CCTTGC CTCTCACCA <u>T</u> CCTTGC
IL-18 -137 G/C (RS187238)	CACAGAGCCCCAACTTTTACG GGCAGAGGATACGAGTACTTCTTTT	ACTATTTTCATGAAAT <u>C</u> TTTT CT TTTTCATGAAAT <u>G</u> TTTTCT
TNFα -238 G/A	CAGTCAGTGGCCAGAAGAC	CTCGGAATC <u>G</u> GAGCAG

SNP	Primers	Probes
(RS361525)	CCCTCACACTCCCCATCCT	CTCGGAATC <u>A</u> GAGCAG
RANTES 1.1 G/A	TGCTTCATGGCAGGGATCTC	TTTTTCTGTCTTT <u>A</u> AAGGTCTA C
(RS2280789)	GTGAACACCTGTAGGCCTTGAG	CTGTCTT <u>C</u> AAGGTCTAC

Electronic version, part 3:

Antibiotics in CAP defined as appropriate or inappropriate therapy according to the most frequently identified causative micro-organism (for further information see reference 31).

	Streptococcus pneumoniae ¹	Haemophilus influenzae ²	Legionella pneumophila	Mycoplasma pneumoniae	Staphylococcus aureus ³
Penicillin	+	–	–	–	–
Amoxicillin	+	+	–	–	–
Amoxicillin- clavulanate	+	+	–	–	+
Cephalosporins	+	+	–	–	+
Macrolides	–	–	+	+	–
Quinolones	–	+	+	–	–

+ = appropriate, – = inappropriate antibiotic therapy

¹ all *S. pneumoniae* penicillin sensitive, ² all *H. influenzae* amoxicillin sensitive, ³ no MRSA