

Supplemental Table 1. Effects of IL-4/IL-13 in GC

Research object	Methods	Main results	Reference
CRL1739 cells, human GC specimens	Immunoprecipitation and immunoblot analysis	IL-4R α and IL-2R γ -common chain were expressed in CRL1739 cells and GC specimens.	42
CRL1739 cells	Protein synthesis assay using [³ H]leucine	One chimeric protein composed of IL-13 and Pseudomonas exotoxin was highly cytotoxic.	39
HTB-135 cells	Cell Cycle Analysis Kit (Becton Dickinson), FCM	IL-4 inhibited proliferation of GC cells by blocking cell cycle progression.	43
MKN-45 and AGS cells	Boyden chamber assays	CHI3L1 secreted by M2 macrophage could promote the metastasis of GC cell lines by binding to the IL-13R α 2 chain.	40
AGS and MGC308 cells; 100 human GC tissues	Co-immunoprecipitation assay, immunofluorescence, IHC	CD44v3 bound to both CHI3L1 and IL-13R α 2 in GC cell lines. CHI3L1 was upregulated during GC development.	32
HTB13, CRL1863, CRL1739, and KATO III cells; one primary GC cell line	Standard [³ H]thymidine incorporation proliferation assay, FCM	IL-4 could inhibit the growth of GC cells and this effect was positively related with IL-4R expression level of the respective cell lines.	44
507 GC patients	Tissue microarray and IHC	Overexpression of IL-13R α 2 chain in GC tissues was associated with poor prognosis after gastrectomy.	15
30 biopsies from GC patients	IHC	The expression of IL-4 was higher in stages I and II GC than in stages III and IV.	46
17 patients and 30 healthy controls	sandwich ELISA kit (Affymetrix eBioscience)	Serum IL-4 levels in patients were significantly higher than in controls.	45
25 patients and 54 healthy controls	Bio-Plex human cytokine assay (Bio-Rad)	Cytokine levels of IL-4 and IL-13 in patients' plasma were significantly higher than in controls.	41

Cell lines mentioned in this table are all human GC cell lines unless otherwise indicated. GC: gastric cancer; IHC: immunohistochemistry; CHI3L1: Chitinase 3-like protein 1; ELISA: enzyme-linked immunosorbent assay.

Supplemental Table 2. Effects of IL-4/IL-13 in CRC cells

Research object	Methods	Results	Reference
LS174T and HT-29 cells	WB, RT-PCR	IL-4 and IL-13 increased mucin 2 expression in LS174T cells, but not in HT-29 cells.	48
HT-29 cells	WB, RT-PCR, Northern blot; L-[¹⁴ C]valine	IL-13 could inhibit the macroautophagy pathway via the activation of the class I PI3K.	49
HT-29 cells	WB; [¹⁴ C]valine	Overexpression of tumor suppressor phosphatase and tensin homolog could counteract the IL-13 down-regulation of macroautophagy.	50
Colo201 and Colo205 cells	Protein synthesis assay using [³ H]leucine	One chimeric protein composed of IL-13 and Pseudomonas exotoxin was highly cytotoxic.	39
HT-29 and WiDr cells	WB	IL-13 induced phosphorylation of JAK2, JAK1, and Tyk2. IL-13 and IL-4 could induce phosphorylation of STAT6.	51
HT-29, CaCo-2, DLD-1, T84 cells	FCM, Northern blot, IP	IL-4 and IL-13 could upregulate expression of CD44.	52
RKO and SW480 cells	MTT, transwell assay, RT-PCR, WB, luciferase assay	Propofol suppressed cell proliferation and IL-13 induced EMT in CRC cell lines. Propofol suppressed IL-13/STAT6 signaling pathway.	54
HT29, SW480, Caco2 cells	WB, PCR, transwell assays, CHIP, luciferase reporter assay	IL-13 promoted EMT and aggressiveness of CRC through IL-13R α 1/STAT6/ZEB1 pathway.	55
HT-29, WiDr, SW1116, Co-115, LS411N, LS513 and LS1034 cells	MTT assay and incorporation of [³ H]-TdR; FACS; northern blot; biotin-labeled IL-4 binding studies	In HT29, WiDr, LS411N, LS513, LS1034 cells, IL-4 inhibited proliferation; in CO-115 and SW1116 cells, no difference. Three cell lines (HT29, LS1034, WiDr) exhibited relatively high density of the IL-4R and 4 (Co-115, LS411N, LS513, SW1116) had lower density.	65
HT-29 and DLD-1 cells	Cell proliferation was determined by counting the cells.	IL-4 and IL-13 increased nicotinamide adenine dinucleotide phosphate oxidase 1-related proliferation.	12
HT-29 and WiDr cells	[³ H]-TdR incorporation assay	IL-4 inhibited cell growth of CRC cells.	69
HT-29 cells	Cell counting or colorimetric MTT assay.	IL-4 inhibited the growth of HT-29 cells.	70
HTB 38 cells	Human tumor cloning assay	IL-4 showed antiproliferative activity.	71
LS531 cells	[³ H]-TdR incorporation assay	IL-4 inhibited cell proliferation.	73
HT-29 cells	FACS. PBMCs were separated from a patient with chronic lymphocytic leukemia.	IL-4 significantly suppressed IL-12, IL-2 and IFN- α enhanced ADCC against HT-29 cells.	75

SW948 cells, human PBMCs, mAb 17-1A	PBMCs were incubated for 0-24 h with IL-4 before ADCC assay	Pretreatment with 1 ng/ml IL-4 for 2 hours induced a significant increase in the ADCC activity of PBMCs.	76
LS174T cells; human peripheral monocytes	24-h ¹¹¹ In-release assay was used to measure ADCC activity.	IL-4 and macrophage colony-stimulating factor had a synergistic enhancement effect of monocyte- mediated ADCC on LS174T cells.	78
HT-29 cells, PBMCs	Flow cytometric cytotoxicity assay	IL-4 decreased ADCC and reduced the IL-2, IL-12, and IFN-alpha-induced ADCC.	79
SW620 cells	ELISA	IL-4 sensitized SW620 cells to radiation by inhibiting NF-kappaB.	80
HT-29, LoVo and SW480 cells	Indirect immunofluorescence, FACS	IL-4 could decrease epithelial cellular adhesion molecule and Lewis ^y expression in HT-29 and LoVo cells, but not in SW480 cells.	81
LS174T cells	Northern blot, RT- PCR, FCM	IL-4 inhibited the expression of stem cell factor and its receptor c-kit in LS174T cells. IL-4 could also inhibit stem cell factor-induced proliferation.	82
HRT18, H29/6 and HT115 cells	ELISA	The addition of IL-4 increased IL-8 release in HT115 cells, but not in HRT18 and H29/6 cells.	83
Colo205 cells	FCM, RT-PCR analysis	IL-4 inhibited cell-cell adhesion but not cell proliferation. IL-4 and IL-13 could down-regulate E- cadherin and carcinoembryonic antigen molecules.	84
Primary CRC cell lines CoSp, Cope, and CoDo; HT-29 cells	WB, RT-PCR, transwell assay	IL-4 was an inhibitor of hepatocyte growth factor and it could regulate hepatocyte growth factor- induced cell proliferation and other events of tumor progression.	85
WiDr and HT- 29 cells	[³ H]-TdR uptake studies, IP and WB, northern analysis	The addition of IL-4 phosphorylated JAK1, JAK2, and Tyk2, and activated JAK1 and JAK2.	86
Caco cells	RT-PCR, MTT assay, trypan blue test, ELISA	Anti-IL-4 antibody inhibited the growth of the Caco cells. Neutralizing of IL-4 increased the efficacy of chemotherapy and inhibited the CD133+ cells.	88
HT-29 and Caco-2 cells	RT-PCR, FCM, transwell migration assay	IL-4/Stat6 activities correlated with apoptosis and metastasis in colon cancer cells.	89

Cell lines mentioned in this table are all human colon cancer cell lines unless otherwise indicated. CRC: colon and rectal cancer; WB: Western blot; RT-PCR: reverse transcription-polymerase chain reaction; PI3K: phosphatidylinositol 3-kinase; JAK: Janus kinase; STAT: signal transducer and activator of transcription; FCM: flow cytometry; MTT: 3-(4,5-Dimethylthiazol-2-yl)-2,5-diphenyltetrazoliumbromid; CHIP: chromatin immunoprecipitation; IP: immunoprecipitation; EMT: epithelial-mesenchymal transition; ZEB1: Zinc finger E-Box binding homeobox 1; FACS: fluorescence-activated cell sorting; IFN: interferon; ADCC: antibody-dependent cellular cytotoxicity; PBMC: peripheral blood mononuclear cell; PKH: Paul Karl Horan; ELISA: enzyme-linked immunosorbent assay; NF-κB: nuclear factor kappa-light-chain-enhancer of activated B-cells; EpCAM: epithelial cellular adhesion molecule.

Supplemental Table 3. Effects of IL-4/IL-13 in CRC mouse models or patients

Research object	Methods	Results	Reference
KM12, KM12C and KM12SM cells, nude mice and 80 CRC patient tissues	ELISA, WB, IHC, FCM	High expression of IL-13R α 2 in KM12 cells was associated with liver metastasis in nude mice. IL-13R α 2 overexpression in CRC tissues was associated with later stages and poor prognosis.	13
Tissues from 359 patients	IHC	High expression of IL-4, IL-4R, and IL-13R was associated with a lower frequency of lymph node metastases.	47
KM12SM, SW620, a metastasis model of nude mice	WB, IHC, IP, transwell assay	IL-13R α 2 induced activation of the FAK and PI3K/AKT/mTOR pathways and this was mediated by FAM120A.	11
SW480 cells and murine CC cells	WB, quantitative RT-PCR, MTT assay, colony formation assay; nude mice	IL-13 induced the expression of 11 β HSD2 in CRC cells through IL-13R α 2 and promoted the malignancy of CRC.	56
CT26, murine liver metastasis model	Mass spectrometry analysis, WB, IP, MTT assays, FCM	Protein tyrosine phosphatase-1B could mediate IL-13-induced cancer cell proliferation, migration, invasion and survival.	57
KM12SM, SW620, RKO cells, mouse models	ELISA, real-time PCR, WB, IHC	IL-4 and IL-13 both up-regulated the expression of chemokine eotaxin-2 in LS174T and LOVO cells.	58
LS174T, LOVO, SW480 and COLO 205 cells; 25 CRC patients	WB, IHC, MTT assay, Bio-Plex Pro Mouse Cytokine 23-Plex Panel	The formation of CRC was more frequent in obese mice than wild type mice. Silencing IL-13R α 1 inhibited IL-13-induced proliferation in HT29.	59
Mouse model of CRC induced by injection of azoxymethane; HT29 cells	FCM, BALB/c mice and CD1-knockout mice	The development of lung metastases could be significantly decreased by an IL-13 inhibitor, but not by inhibiting IL-4.	60
Murine CRC lung metastasis model; murine CC cell line CT26	Competition ELISA, MTT assay, wound healing assay, transwell invasion assay, FCM	IL-13R α 2 D1 peptide inhibited migration, invasion, and proliferation in metastatic CRC cells treated with IL-13. Nude mice treated with the enantiomer D-D1 peptide showed a significantly survival increase to metastasis.	61
KM12SM and SW620 cells, nude mice and Balb/c mice	IHC, case-control epidemiology study, FCM, multiplex immunoassay	Reduced IL-4R signalling increased CRC risk but reduced tumor progression. It had no effect on CRC mortality.	62
IL-4R α and IL-13 transgenic mouse models, CRC patients and controls	ELISA	Patients with advanced stage of CRC had lower serum IL-13 levels. Low serum IL-13 was associated with poorer prognosis.	63
241 patients with CRC			

Fecal protein from 20 CRC patients and 20 healthy controls	ELISA, bicinchoninic acid assay	The expression of IL-13 was significantly higher in CRC patients.	64
Murine CC cell line CT26 tumor-bearing mice	ELISA, quantitative RT-PCR	Protein levels of IL-4, and IL-13 were higher in the serum or the tumor homogenates of CT26 tumor-bearing mice.	66
Patients with metaplastic polyps, adenomas, or carcinomas	IL-4R was detected by IHC	The expression of IL-4R was positive in all polyps (5/5), adenomas (15/15) and in 40/45 carcinomas.	67
HCT116, HT-29, DLD-1, SW480, SW620, Caco2 and HCA7 cells; IL-4R α -/- mice	MTT assay, RT-PCR, WB, FCM, immunostaining	IL-4R α expression promoted tumor growth in HCT116, HT-29, DLD-1, SW480, SW620, Caco2 and HCA7 cells, but IL-4 could only decrease apoptosis in HCT116 cells.	68
HCT116 and RKO cells; subcutaneous xenograft mouse models, 218 human CRC samples	WB, qRT-PCR, luciferase assay, chromatin IP assay, transwell invasion assay, IHC	IL-4 could promote EMT of HCT116 and RKO cells through E2F1/SP3/STAT6 axis. Analysis of clinical CRC samples showed a positive correlation between E2F1, SP3 and STAT6.	74
HT-29 cells and a metastasis model in nude mice	FCM, northern blot	IL-4 changed the expression of integrin and decreased the lung-colonizing ability of HT-29 cells.	87
Primary human CC cells; nude mice	FCM, IHC, immunofluorescence analysis, WB	CC growth was dictated by stem-like cells which were resistant to chemotherapy due to autocrine of IL-4.	14
CC-bearing mice	qRT-PCR, ELISA, immunofluorescence	Addition of IL-4 improved muscle function and lifespan of CC-bearing mice.	90
HCT116 cells; NOD/SCID mice; CC tissues from 40 patients	RT-PCR, IHC, WB, FCM, transwell migration assay	Over-expression of IL-12 could inhibit the expression of IL-4 and STAT6 in CSCs and inhibit their survival.	91
MC38 murine CC cell line, mice	IHC, 4-h 51Cr-release assays, female C57BL/6 (B6) mice	IL-4 combined with CpG oligonucleotide could suppress tumor growth by activating Th1-type immune responses.	92
Murine CC cell line CT26 tumor-bearing mice	IHC, FCM, qRT-PCR	An IL-4R α aptamer-liposome-CpG oligodeoxynucleotide delivery system had enhanced anti-tumor activity.	93
Murine CC cell line colon 26; BALB/c mice	IL-4 overexpressing colon 26 cells; ELISA; 51Cr release assays	Overexpression of IL-4 in colon 26 cells could induce local tumor killing as well as systemic immunity in mice.	94
MC38 murine CC cell line, female B6 mice	Subcutaneous tumor models of mice; 4-h 51Cr-release assays	IL-4 overexpression MC38 cells promoted a tumor-specific Th1-type response in B6 mice.	95

66 patients with CRC and 87 healthy controls	multiplex bead array immunoassay (Millipore)	Serum levels of IL-4 were higher in CRC patients.	96
77 CRC patients, 70 healthy controls	ELISA	Serum IL-4 levels of the patients were significantly higher than the control group.	97
105 CRC patients	ELISA	The serum levels of IL-4 can be used as a marker of pre-invasive to invasive CRC.	98
10 CRC patients with liver metastases and 5 healthy controls	Flow cytometric immunophenotyping	Higher serum levels of IL-4 were found in patients than controls. The CD8+ cytotoxic cells might be negatively regulated by serum IL-4.	99
99 CRC patients and 107 healthy controls	ELISA	IL-4 serum levels were significantly higher in patients.	100
20 CRC patients	fluorescent bead-based detection assay, Multiplex kit	No IL-4 expression was detected in serum, normal mucosa or tumor tissue.	101
Primary CC cells isolated from 18 patients; nude mice	RT-PCR, WB, FCM, transwell assay, FCM, TUNEL staining	IL-4 could increase the expression of survivin by activating STAT6.	102
Primary human CC cells and T84 cells	FCM, IHC, Real-time PCR analysis	Tumor-cell-derived IL-4 enhanced apoptosis resistance in primary human CC cells and T84 cells.	103
Primary and metastatic tumors from four CRC patients	MagSweeper, whole-transcriptome analysis with RNA-Seq	The expression of IL-4 was significantly up-regulated in CD133(+) cells compared to CD133(-) CRC cells	104
Primary CRC cells, NOD/SCID CB-17 mice	Immunofluorescence analysis, ELISA, subcutaneous tumor model of mice	Cancer-initiating cells associated IL-4 was responsible for weak immunogenicity <i>in vitro</i> .	105
49 primary CC and 20 metastases	IHC	Expression of IL-4 by > or = 20% of CC tumor infiltrating lymphocytes was associated with better prognosis.	106
41 metastatic CRC patients	Bio-Plex human cytokine multiplex kits (Bio-Rad Inc.), FACS, IHC	The patients with higher serum IL-4 levels have longer overall survival when receiving the thymidylate synthase poly-epitope-peptide anticancer vaccine.	107
80 CRC patients and 38 matched controls	Human Th1/Th2 Cytokine Kit II (BD Biosciences)	IL-4 was measured from the supernatants of activated peripheral blood mononuclear cells. No significant difference was found.	108
Patients with CRC	ELISA, EFA ingestion	Long-term EFA ingestion could reduce total serum IL-4 by 69% after 6 months.	109

Cell lines mentioned in this table are all human CC cell lines unless otherwise indicated. CRC: colon and rectal cancer; CC: colon cancer; FCM: flow cytometry; ELISA: enzyme-linked immunosorbent assay; WB: Western blot; IHC: immunohistochemistry; FAM120A: family with sequence similarity 120A; RT-PCR:

reverse transcription-polymerase chain reaction; MTT: 3-(4,5-Dimethylthiazol-2-yl)- 2,5-diphenyltetrazoliumbromid; 11 β HSD2: 11 β -hydroxysteroid dehydrogenase type II; IP: immunoprecipitation; qRT-PCR: quantitative real-time PCR; NOD/SCID: nonobese diabetic/severe combined immunodeficiency; CSC: cancer stem cell; FACS: fluorescence-activated cell sorting; Th: T helper; EFA: essential fatty acids.

Supplemental Table 4. Polymorphisms of IL-4/IL-13 and their receptors in GC

First author (year)	Country	Ethnicity	Number of cases/controls	SNPs	Result
El-Omar (2003) (116)	USA	Mixed	112/209	IL-4 (-590 C/T), rs2243250	No significant association
Wu (2003) (117)	China	Asian	220/230	IL-4, rs2243250; IL-4R, rs1805010; IL-4R, rs1801275	Higher risk of developing diffuse type or cardia type cancer was observed for the CT/CC genotype of IL-4 at the position -590 (rs2243250).
Lai (2005) (118)	China	Asian	123/162	IL-4 (-590), rs2243250	No significant association
Garcia-Gonzalez (2007) (119)	Spain	Caucasian	404/404	IL-4 (-590 C>T), rs2243250	No significant association
Crusius (2008) (120)	10 European countries	Caucasian	235-244/1107-1160	IL-4, rs2243250; IL-4, rs2070874; IL-4R, rs1805010; IL-4R, rs2057768	Rs2057768 is related with non-cardia gastric adenocarcinoma.
Zambon (2008) (121)	Italy	Caucasian	144/171	IL-4 -588C>T, rs2243250; IL-4RA Ex5+14A>G, rs1805010; IL-4RA Ex11+828A>G, rs 1801275	No significant association
Ando (2009) (122)	Japan	Asian	330/190	IL-4 -590C/T; IL-4R α Ile50Val	No significant association
Ko (2009) (123)	Korea	Asian	78-81/320-324	rs2070874; rs2243250	No significant association
Wu (2009) (124)	China	Han Chinese population	1042/1099	rs2070874	Compared with the IL-4 -168TT genotype, heterozygous -168TC and combined -168TC/CC genotypes were associated with a significantly decreased GC risk.
Pan (2014) (125)	China	Asian	308/307	rs2243250	No significant association

Yin (2015) (126)	China	Asian	234/465	IL-13, rs1800925	No significant association with gastric cardiac adenocarcinoma risk
Cárdenas (2018) (45)	Colombia	Mixed	15-17/20-30	IL-4 -590 (C/T), IL-4 Ile50Val, IL-4 Q576R	No significant association
Martínez-Campos C (2019) (127)	Mexico	Mixed	124/125	IL-4-590C/T (rs2243250)	No significant association
He B (2019) (128)	China	Asian	479/483	IL-4 rs2243248; IL-4 rs2070874; IL-4R rs2057768; IL-4R rs2107356; IL-4R rs1805015; IL-4R rs1801275	No significant association
Yun Y (2017) (129)	China	Asian	340/364	IL-4 rs2243250 (590 C/T), rs2227284 (107 T/C), rs2070874 (168 T/C) and rs1801275 (576 Q/R)	The IL-4 rs2243250 CC genotype and CT+CC genotype were associated with higher GC risk.
Burada F (2012) (130)	Romania	Caucasian	105/242	IL-4R -3223C→T, rs2057768	IL-4R -3223C→T polymorphism may increase the risk of gastric adenocarcinoma, mainly for the noncardia type.
Xia HZ (2012) (131)	China	Asian	308/425	IL-4R, rs2107356(G>A)	A significant reduction of the IL-4R AA genotype in GC risk was found when compared to GG genotype.
Bhayal AC (2015) (132)	India	Asian	182/326	IL-4 intron 3 variable number of tandem repeat (VNTR)	The study revealed an association of 2R allele and 2R carrier genotypes in the etiopathogenesis of GC.
Sampaio AM (2015) (133)	Portugal	Portuguese	100/50	IL-4 (-1098T>G), rs2243248; IL-4 (-590C>T), rs2243250; IL-4 (-33C>T), rs2070874	IL-4-590TT and IL-4-1098GG were found associated with intestinal type GC and diffuse type GC, respectively. IL-4 TTT haplotype was linked with both intestinal and diffuse type GC groups.
Seno H (2007) (134)	Japan	Asian	100/93	IL-4 (11 SNPs), IL-4R (22 SNPs)	The IL-4 gene diplotypes are negatively associated with the risk of developing GC due to Helicobacter pylori infection.

Wang Y (2017) (135)	China	Asian	362/384	IL-4, rs2243248 (-1098 G/T), rs2227284 (-33 C/T), rs2243250 (-589 T/C) and rs2070874 (-107 T/C)	The TC and CC genotypes of rs2243250 were associated with an increased risk of GC. The TTTT haplotype revealed a reduced risk of GC.
Pavithra D (2018) (136)	India	Asian	200/400	IL-4C-590-T	No significant association
Cavalcante (2017) (137)	Brazil	Mixed	119/474	IL-4, rs79071878, VNTR	rs79071878 was positively associated with the development of GC.
Sarah Yang (2017) (138)	Korea	Asian	368-377/736-754	IL-4R (rs7205663 and rs1805010); IL-13 (rs6596090, rs20541)	No significant association
Wang YM (2016) (139)	China	Asian	132/1198	IL-4R rs2057768	No significant association
Schmidt HM (2011) (140)	Singapore	Asian	60/162	IL-13, rs1800925; IL-4, rs2070874; IL-4R, rs1801275.	No significant association
García-González MA (2012) (141)	Spain	Caucasian	380/-	IL-4, rs2243250	Not relevant in determining the prognosis of gastric adenocarcinoma
Talebkhani Y (2017) (142)	Iran	Caucasian	31/46	IL-4 C-590T	Serum antibodies against Helicobacter pylori neutrophil activating protein in carriers of IL-4 C-590T genetic polymorphism amplify the risk of GC.

GC: gastric cancer; SNPs: single nucleotide polymorphisms.

Supplemental Table 5. Polymorphisms of IL-4/IL-13 and their receptors in colorectal cancer (CRC)

First author (year)	Country	Ethnicity	Number of cases/controls	SNPs	Result
Walczak A (2011) (143)	Poland	Caucasian	150/170	IL-13, rs1800925	The CT genotype is connected with a higher risk of colon cancer occurrence.
Ibrahimi M (2019) (144)	Iran	Iranian	123/152	IL-4 VNTR, RP1/RP1, RP1/RP2	No significant association
Landi S (2007) (145)	Spain	Caucasian	377/326	IL-4 (-588C>T, Ex1-168G>A); IL-4R (I75V, C431R, S436L, S503P, Q576R)	The homozygotes for IL-4 -588C>T or for Ex1-168G>A showed an increased risk for colon cancer. Women showed an increased risk associated to the IL-4 rare alleles.
Lee YS (2009) (146)	Korea	Asian	170/130	IL-4R 1902	The IL-4R 1902*T allele was found to be associated with an increased risk of colon cancer and rectal cancer, while IL-4R 1902*C allele was associated with a decreased risk.
Cavalcante GC (2017) (137)	Brazil	Mixed	63/474	IL-4, rs79071878, VNTR	No significant association
Jose M Cozar (2007) (147)	Spain	Caucasian	96/176	IL-4-590 C>T (rs2243250)	No significant association
Suchy (2008) (148)	Poland	Caucasian	607/350	IL-4 (-590 C/T)	No significant association
Wilkening (2008) (149)	Sweden	Caucasian	308/585	IL-4-590C/T (rs2243250); IL-4R-3223G/A (rs2057768)	The rare T allele of IL-4-590 was related to a longer survival. The prognostic value of the genotypes was of borderline significance.

Levar Shamoun (2018) (150)	Sweden	Caucasian	466/445	IL-4, rs2243250; IL-4R α , rs1801275; IL-13, rs1800925	IL-13 SNP rs1800925 is a risk factor for CRC and that IL-4 SNP rs2243250 could be a useful prognostic marker.
Yanming Yu (2017) (151)	China	Asian	513/572	IL-13: rs847 A>G; IL-13: rs848 T>G; IL-13: rs1295685 C>T	A significant antagonistic interaction was found between rs848 (G-T) and allium vegetable intake; moreover, significant combined and synergistic interactions were observed for all three SNPs and overnight meal intake.
Y Yu (2014) (152)	China	Asian	299/296	IL-4 rs2070874	No significant association
Diego Marques (2017) (153)	Brazil	Mixed	140/140	IL-4 rs79071878	Polymorphism variations in IL-4 gene was associated with increased CRC risk.
Walczak A (2012) (154)	Poland	Caucasian	191/205	IL-13 - 1112 C/T, rs1800925	The CT and TT genotypes of the IL-13 - 1112 C/T polymorphism may be connected with a higher risk of CRC.
Raul Zamora-Ros (2015) (155)	Spain	Caucasian	274/266	IL-4, rs2243250	The novel dietary inflammatory index (DII) score was inversely correlated with SNP rs2243250, and an interaction was observed with CRC risk.
Xiulin Wen (2020) (156)	China	Asian	248/463	IL-4, rs2243250 and rs2227284	rs2243250 and rs2227284 in IL-4 are associated significantly with reduced CRC risk.
Nicola Ingram (2013) (62)	United Kingdom	Caucasian	1502/584	IL-4R α : rs1801275, rs1805015, rs1805016, rs1805013, rs1805011, rs1805010.	SNP rs1801275 was associated with increased CRC risk. Reduced IL-4R signalling was associated with increased CRC initiation and risk but reduced tumour progression and no effect on CRC mortality.
Florin Burada (2013) (157)	Romania	Caucasian	144/233	IL-4R -3223C > T	No significant association
Yannopoulos (2007) (158)	Greece	Caucasian	93/108	IL-4 (-590 C/T)	The (-590 C/T) polymorphism in the IL-4 gene is associated with increased risk for early stages of colorectal adenocarcinoma.

Bente A Talseth (2007) (159)	Poland	Caucasian	118/100	IL-4 C-589T (rs2243250)	No significant association between SNP rs2243250 and hereditary non-polyposis CRC.
Juan Sainz (2012) (160)	Germany	Caucasian	1798/1810	IL-13, rs20541	Patients harboring the IL-13_rs20541_T allele had a reduced risk of CRC.
Lin Xiao (2016) (161)	China	Asian	58/-	IL-13, 1112 C/T (rs1800925)	The studied SNP does not predict responsiveness to neoadjuvant chemoradiotherapy or prognosis of locally advanced rectal cancer.
A Ho-Pun-Cheung (2011) (162)	France	Caucasian	71/-	IL-4, rs2243250; IL-13, rs20541, rs1800925	The SNP IL-13 rs1800925 was significantly associated with rectal adenocarcinoma response to chemoradiation.

CRC: colon and rectal cancer; SNPs: single nucleotide polymorphisms.