

Supplementary Table 1 Disease characteristics of the patients with Crohn's disease enrolled in the study

Paris Classification		At diagnosis n (%)	At worst flare n (%)
Age	A1a: 0-10 years	53 (24.8)	21 (9.8)
	A1b: 10-17 years	148 (69.2)	142 (66.4)
	A2: 10-40 years	12 (5.6)	21 (9.8)
Location	L1: Ileal	52 (24.3)	40 (18.7)
	L2: Colonic	39 (18.3)	27 (12.6)
	L3: Ileocolonic	98 (45.8)	93 (43.7)
Behaviour	L4a: upper disease proximal to ligament of Treitz	22 (10.3)	18 (8.4)
	L4b: upper disease distal to ligament of Treitz	8 (3.7)	9 (4.2)
Growth	B1: nonstrictureing	148 (69.5)	116 (54.5)
	B2: structuring	15 (7.0)	19 (8.9)
	B3: penetrating	19 (8.9)	21 (9.8)
	B2B3: penetrating and stricturing	4 (1.9)	5 (2.3)
	P: perianal disease modifier	20 (9.3)	20 (9.3)
	G0: no evidence of growth delay	147 (69.0)	124 (58.2)
	G1: growth delay	33 (15.4)	34 (15.9)

Supplementary Table 2 Disease characteristics of the patients with ulcerative colitis enrolled in the study

Paris Classification		At diagnosis n (%)	At flare n (%)	worst
Extent	E1: Proctitis	19 (9.9)	9 (4.7)	
	E2: Left-sided colitis	33 (17.2)	28 (14.6)	
	E3: Extensive colitis	30 (15.6)	24 (12.5)	
	E4: Pancolitis	90 (46.9)	82 (42.7)	
Severity	S0: never severe	116 (60.4)	89 (46.4)	
	S1: ever severe	40 (20.8)	52 (27.1)	

Supplementary Table 3 Genotype distribution in the Polish population according to previous studies

	Study group (n)	$\epsilon 2\epsilon 2$	$\epsilon 2\epsilon 3$	$\epsilon 3\epsilon 3$	$\epsilon 3\epsilon 4$	$\epsilon 4\epsilon 4$	$\epsilon 2\epsilon 4$
The current study	IBD (406)	3	53	253	88	4	5
Kowalska et al. ⁴¹	Healthy subjects (137)	0	11	100	19	3	4
Bednarska-Makaruk et al. ⁴²	Healthy subjects (170)	2	18	115	30	1	4
Jasieńska et al. ⁴⁰	Healthy women (118)	0	10	89	15	3	1
Bojar et al. ⁴³	Postmenopausal women (402)	nd	63	253	*		

*only data for APOE ϵ 4 allele were available: 86

Supplementary Table 4 Clinical characteristics in patients with inflammatory bowel diseases depending on major APOE genotypes

Variables	n	$\epsilon 2/\epsilon 3$	$\epsilon 3/\epsilon 3$	$\epsilon 3/\epsilon 4$	p value

Median (IQR) or n (%)						
Age [y]						
at inclusion	39	15.6 (13.2-16.8)	15.2 (13.1-17.1)	14.5 (12.4-16.3)	0.236	
	7				4	
at diagnosis	40	11.7 (9.3-14.6)	12.7 (8.9-14.8)	12.5 (9.2-14.3)	0.818	
	4				2	
at worst flare	35	13.6 (11.5-15.9)	13.6 (10.8-15.9)	13.9 (10.7-15.8)	0.992	
	5				1	
Duration of the disease [y]	39	2.8 (0.9-5.4)	2.0 (0.6-3.9)	1.7 (0.4-4.1)	0.188	
	0				0	
Nutritional status						
Weight at diagnosis [kg]	38	38.3 (28.2-49.5)	38.0 (25.5-52.9)	43.3 (28.4-54.3)	0.560	
	7				2	
Weight diagnosis [z score]	38	-0.43 (-1.01-0.28)	-0.73 (-1.37-0.00)	-0.56 (-1.23-0.41)	0.079	
	3				1	
Height at diagnosis [cm]	38	148.3 (138.3-167.0)	152.0 (134.0-166.0)	152.5 (138.5-169.3)	0.807	
	2				1	
Height diagnosis [z score]	37	-0.15 (-0.82-0.61)	-0.28 (-1.15-0.53)	0.12 (-0.84-1.00)	0.062	
	8				2	
Body mass index at diagnosis [kg/m ²]	38	17.24 (15.65-18.87)	16.82 (14.80-18.96)	17.18 (15.17-20.52)	0.507	
	2				2	
Body mass index at diagnosis [z score]	37	-0.30 (-1.18-0.11)	-0.67 (-1.28-0.03)	-0.59 (-1.19-0.51)	0.438	
	8				9	
Weight at worst flare [kg]	34	43.3 (34.0-53.0)	43.0 (30.5-54.5)	48.5 (34.5-59.5)	0.358	
	5				3	

Weight at worst flare [z score]	33	-0.55 9	([-1.06]- 0.23)	-0.81 0.05])	([-1.40]-[- 0.35])	-0.55 0.35)	([-1.04]- 4)	0.080
Height at worst flare [cm]	34	153.3 5	(146.5- 168.0)	158.0 168.8)	(141.0- 172.0)	160.0 172.0)	(144.5- 3)	0.627
Height at worst flare [z score]	34	0.01 0	([-0.86]- 0.75)	-0.36 0.49)	([-1.13]- 0.81)	-0.17 0.81)	([-0.91]- 2)	0.187
Body mass index at worst flare [kg/m ²]	34	17.64 1	(15.86- 19.45)	17.10 19.53)	(15.24- 21.29)	17.87 21.29)	(15.79- 1)	0.264
Body mass index at worst flare [z score]	33	-0.66 7	([-1.16]- 0.07)	-0.80 0.11])	([-1.36]-[- 0.54])	-0.48 0.54)	([-1.22]- 4)	0.251

Albumin level

at diagnosis [g/dl]	34	4.1 (3.8-4.4) 5	4.0 (3.5-4.3)	4.0 (3.6-4.4)	0.263 7
at worst flare [g/dl]	32	4.0 (3.8-4.5) 7	3.9 (3.4-4.3)	4.1 (3.8-4.4)	0.033 5

Parameter of

inflammation

CRP at diagnosis [mg/l]	38	4.0 (0.8-22.8) 6	6.6 (1.2-20.9)	3.9 (0.5-18.3)	0.249 0
CRP at worst flare [mg/l]	34	4.7 (1.4-28.8) 7	7.7 (1.9-31.3)	3.0 (0.5-17.2)	0.016 5

Treatment

Systemic steroids ¹	40	30 (56.6) 6	165 (65.2)	50 (56.8)	0.250 4
Number of courses of steroid treatment	40	1 (0-2) 2	1 (1-2)	1 (0-2)	0.117 5
Immunosuppressive treatment ²	40	34 (64.2) 5	184 (73.0)	56 (63.6)	0.163 8

Number of immunosuppressants	40	1 (0-1)	1 (0-1)	1 (0-1)	0.097
	5				6
Time-to-first dose of immunosuppressive treatment [months]	27	2.7 (0.0-17.0)	2.5 (0.0-9.1)	1.9 (0.0-9.6)	0.556
	5				7
Age at first intake of immunosuppressive treatment [years]	40	13.0 (10.3-14.9)	12.9 (9.4-14.6)	12.0 (9.2-15.5)	0.677
	5				5
Biological therapy ³	40	19 (35.8)	102 (40.3)	31 (35.2)	0.635
	6				7
Total number of biologics	40	0 (0-1)	0 (0-1)	0 (0-1)	0.513
	6				6
Time-to-first dose of biological treatment [months]	15	17.8 (10.0-44.0)	13.9 (7.0-26.8)	12.2 (4.7-26.9)	0.406
	0				2
Age at first biological treatment	15	13.9 (13.5-15.7)	13.1 (10.7-15.1)	13.4 (10.1-15.6)	0.354
	1				9
Operative treatment ⁴	40	2 (3.8)	22 (8.7)	9 (10.2)	0.388
	6				8
Age at first surgery [years]	36	9.5 (7.7-11.4)	14.5 (11.7-16.7)	14.9 (14.0-15.6)	0.037
	8				8
Time-to-first surgery [months]	30	12.0	16.7 (3.0-38.6)	25.1 (7.5-40.5)	0.825
	4				4

Hospitalizations (if duration ≥1 years)

Hospitalizations for relapse (per 1 year of the disease)	0.35 (0.19-0.69)	0.57 (0.28-0.93)	0.61 (0.25-1.28)	0.159
Days of hospitalization for relapse (per 1 year of the disease)	2.46 (0.57-5.11)	4.74 (1.63-8.34)	6.03 (1.66-8.68)	0.074
Relapses from diagnosis (per 1 year of the disease)	0.37 (0.16-0.87)	0.57 (0.28-0.99)	0.60 (0.23-1.33)	0.354
Severe relapses from diagnosis (per 1 year of the disease)	0.00 (0.00-0.29)	0.17 (0.00-0.53)	0.25 (0.00-0.47)	0.176
Concomitant diseases⁵	40	25 (47.2)	81 (32.0)	23 (6.1)
Extraintestinal manifestations	40	10 (18.9)	57 (22.5)	21 (23.9)
	6			2

¹ Systemic steroid therapy included: methylprednisolone, prednisone, hydrocortisone.

² Immunosuppressive and anti-inflammatory agents included: azathioprine, methotrexate, mercaptopurine, cyclosporine, mycophenolate mofetil, tacrolimus, sulfasalazine.

³ Biological agents included: infliximab, adalimumab, golimumab, vedolizumab.

⁴ Only surgery related to IBD-specific problems (e. g. colectomy, resection, fistula, perforation, abscess) was included.

⁵ e.g. celiac disease, bronchial asthma, obesity, gastroesophageal reflux disease, epilepsy, hypothyroidism.