Supplemental Information

Spatiotemporal analysis of human intestinal

development at single-cell resolution

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Table S1: Overview of key genes from scRNA-seq data, divided by each cell compartment, that exhibit relatively high or specific expression in each cell types used for characterisation

annotation, related to Figure 1 (afurther sub-division of secretory cells is also shown, bComparison made as proportion of each compartment cells, if cluster is locational specific comparison of time-point difference made only in that location and overall. Locational and time-point differences: Wilcox rank test, p-value <0.05 *; p-value <0.01 ***; p-value <0.01****)

Compartment	Compartment Genes	Cell Type	Key Genes (Cell Type)	Cluster Name (Graph abstraction number)	Key Genes (Cluster)	Locational Differences	Time-point Differences ^b	
щр		Stem	LGR5, ASCL2, MYC	Proximal Stem Cells (1)	CASP1, GRIA4, PLA2G2A	` '	No significant differences	
				Distal Stem Cells (2)	LEFTY1, PTGDR, SMOC2	Increased in Colon (****)	Increases over time in colon and overall	
		TA	MKI67, TOP2A, UBE2C	Proximal TA (3)	CENPM, TK1, UBE2T	Increased in small intestine (****)	Decreases over time overall	
				Distal TA (4)	HMGB3, MND1, TACC3	Increased in colon (****)	No significant differences	
		Secretory ^a	FOXA2, INSM1, HES6	Secretory Progenitors (5-15)	ISL1, NEUROG3, GHRL	Increased in small intestine (**)	No significant differences	
				EECs (5-15)	CHGA, TPH1, NEUROD1	No locational preference	Increases over time in colon, TI and overall	
	EPC,			Goblet cells (5-15)	MUC2, SPINK4, WFDC2	No locational preference	Increases over time in colon and overall	
Epithelial	EPCAM, FABP1			Proximal Progenitors (16)	FGG, SOX11, FGB	Increased in small intestine (**)	Decreases over time in TI and overall	
lial	IBP1		FABP2, APOE (proximal), FAM3D (distal)	Proximal Early Enterocytes (17)	TF, AFP, VTN	Increased in small intestine (**)	No significant differences	
				Proximal Enterocytes (18)	CCL25, HEBP1, OSR2	Increased in small intestine (***)	No significant differences	
		Absorptive		Proximal Mature Enterocytes (19)	APOC3, ACE, FABP2	Increased in small intestine (**)	Increases over time in TI and overall	
		Absorptive		Distal Absorptive (20)	CA12, HMGCS2, SLC13A2	Increased in colon (****)	No significant differences	
				Distal Enterocytes (21)	SLC26A2, FABP1, CKB	Increased in colon (****)	No significant differences	
				Distal Mature Enterocytes (22)	AQP8, FABP6, IL32	overall		
			N/A	BEST4/OTOP2 cells (23)	BEST4, OTOP2, CA7	No locational preference	Increases over time in TI	
	N/A	Progenitor	N/A	NEUROG3+ Progenitor (5)	NEUROG3, CLPS, MDK			
		Goblet	MUC2, TFF3	Goblet 1(6)	UBE2C, PBK, MKI67	N/A		
				Goblet 2 (7)	TRABD2A, PTGDR, MYC			
				Goblet 3 (8)	SPDEF, WFDC2, ATOH1			
Sec		EEC	Cluster specific	A-Cells/M-Cells (9)	CLEC4G, MLN, GHRL			
Secretory				D-Cells (10)	SST, HHEX, PCP4			
orya				Enterochromaffin Cells (11)	TPH1, RXFP4, CHGA			
				I-Cells (12)	CCK, GC, TDO2			
				L-Cells (13)	PYY, GCG, UCN3			
				N-Cells (14)	NTS, HOPX, MEP1A			
		Paneth	N/A	Paneth (15)	PRSS2, DEFA5, DEFA6			
	THY1, COL1A2, VIM	Cycling	N/A	Fibroblast G2M phase (24)	CENPF, TOP2A, MKI67	No locational preference	Decreases over time in colon, TI and overall	
			N/A	Fibroblast S-Phase (25)	PCLAF, TYMS, GINS2	No locational preference	Decreases over time in colon, TI and overall	
		Progenitor	N/A	Fibroblast progenitor (26)	HMGA2, MBP, THY1(low)	No locational preference	Decreases over time in colon, TI and overall	
Fibr		S1		S1 (27)	PDLIM3, COL28A1, TFPI2	Increased in colon (**)	Increases over time in colon and overall	
Fibroblast				S1 COL6A5+ (28)	COL6A5, LXN, WNT4	Increased in small intestine (***)	Increases over time in TI	
				S1 <i>IFIT3</i> + (29)	IFIT3, CLEC2B, GPX3	No locational preference	Increases over time in colon, TI and overall	
		S2	FRZB, SOX6, F3	Proximal S2 1 (30)	NPY, CXCL10, BMP2	Increased in small intestine (****)	Increases over time in TI	
				Proximal S2 2 (31)	MMP11, INSC, BMP3	Increased in small intestine (****)	No significant differences	
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				Distal S2 (32)	POSTN, ALKAL2, WNT5B	Increased in colon (****)	Increases over time in colon and
				S3 progenitor (33)	OSR2, NR2F1, PTN(low)	No locational preference	overall Increases then decreases over
				S3 Transitional (34)	GREM1, GDF10, FBLN1	No locational preference	time in colon and overall Increases over time in colon, TI
		00	C7, ASPN, OGN	S3 (35)	RAMP1, SCN7A, SCUBE2	No locational preference	and overall Increases over time in colon, TI
		S3	CI, AGFN, OGN	` '		Increased in colon (*)	and overall Increases over time in colon and
				S3 EBF+ (36) S3 HAND1+ (37)	PRRX1, EBF3, EBF2	Increased in colon(*)	overall Increases over time in colon and
				, ,	COL9A3, DLK1, HAND1	Increased in small intestine (*)	overall Increases over time in TI and
		S4	CCL19, C3, CD74	S4 CXL13+ (38)	CXCL13, TNFSF11, CD24	No locational preference	overall Increases over time in colon, TI
				S4 CCL21+ (39)	CCL21, OLFM3, PTGDS	No locational preference	and overall No significant differences
Myofibroblast /Mesothelium	VIM, FOXF1, TAGLN, RSPO2, RGS5(-)	Cycling	N/A	Myofibroblast G2M phase (40)	MKI67, CENPF, TOP2A	No locational preference	Increases over time overall
			N/A	Myofibroblast S-phase (41)	PCLAF, TYMS, GINS2	No locational preference	Decreases over time in colon
		Myofibroblast	THY1, NKX2-3, ACTG2	Myofibroblast progenitors(42)	ADAMDEC1, MFAP4, ADAM28	No locational preference	and overall Increases over time in colon, TI
				Myofibroblasts (43)	ACTA2, MYH11, GREM2	·	and overall
ast	RSP02	Mesothelium	MSLN, WT1, RSPO1	Mesothelium IL18+(44)	IL18, GATA6, WNT10A	No locational preference	No significant differences
	2,			Mesothelium SOX6+ (45)	SOX6, HPGD, STX2	Increased in colon (**)	No significant differences
		Cycling	N/A	Endothelium G2M-phase (46)	MKI67, CENPF, TOP2A	No locational preference	Decreases over time in colon, TI and overall
			N/A	Endothelium S-phase (47)	PCLAF, TYMS, GINS2	No locational preference	Decreases over time in colon and overall
			GJA4, IGFBP3,, UNC5B	Arterial (L) (48)	HEY1, GJA5, ITGB4	No locational preference	Increases over time in colon and overall
		Autodol		Arterial (M) (49)	UNC5B, DLL4, PRND	No locational preference	Increases over time in colon
E	PECA	Arterial		Arterial (CP) (50)	VWA1, AFG1L, EDNRB	No locational preference	Increases over time in colon, TI and overall
dot	PECAM1, CDH5, CLDN5			Arterial (CP) 2 (51)	CHST1, LGALS1, IGFBP3	No locational preference	Decreases over time in colon and overall
Endothelium		Venous	MADCAM1, APLNR, PRCP	Venous (L) (52)	ACKR1, VWF, ADGRG6	No locational preference	Increases over time in colon and overall
ğ				Venous (M) 1 (53)	LRG1, PLVAP, STMN1(-)	No locational preference	Increases over time in colon, TI and overall
				Venous (M) 2 (54)	CALCRL, APLNR, PRCP	No locational preference	Decreases over time overall
				Venous (CP) 1 (55)	CCL2, CD69, RGCC	No locational preference	Decreases over time in colon and overall
				Venous (CP) 2 (56)	LDHB, JUNB(-), ZFP36(-)	No locational preference	Decreases over time in colon, TI and overall
		Lymphatic	N/A	Lymphatic (57)	LYVE1, PROX1, RELN	No locational preference	Increases over time in colon and overall
	KCNJ8, ABCC9, RGS5, CSP4G	Cycling	N/A	Pericyte G2M-phase (58)	MKI67, CENPF, TOP2A	No locational preference	Decreases over time in TI and overall
			N/A	Pericyte S-phase (59)	PCLAF, TYMS, GINS2	No locational preference	Decreases over time in colon and overall
_			See compartment genes	Pericyte progenitors (60)	THBS1, MBP,CEBPD	No locational preference	Decreases over time in colon and overall
erio		Pericytes		Undifferentiated pericytes (61)	CYGB, NFKBIA, TESC	No locational preference	Increases over time in colon and overall
Pericytes				WNT6+ pericytes (62)	WNT6, GPAT2, THBS1(-)	No locational preference	Increases over time in colon and overall
S				Angiogenic pericyte (63)	PROCR, EBF2, THBS4	Increased in colon (**)	Increases over time in colon and overall
				BMPER+ pericytes (64)	BMPER, CYGB, SPON2	No locational preference	Increases over time in colon, TI and overall
				Contractile pericyte (65)	ADIRF, MYH11, ACTA2	No locational preference	Increases over time in colon and overall
	PHOX2B, HAND2, TUBB2B	Cycling	N/A	Neural G2M & S-Phase (66)	MKI67, CENPF, TOP2A	No locational preference	Decreases in time in colon and overall
Neural		Glial	S100B, SOX10, PLP1	Lymphoid associated glial (67)	FGL2, GFRA3, RXRG	Increased in colon (**)	Increases over time in colon and overall
				Intraganglionic glial (68)	ENTPD2, SPRY1, SOX6	No locational preference	Increases over time in colon, TI and overall
				Submucosal glia (69)	CEBPD, SOCS3, ZFP36	No locational preference	Increases in time in colon and overall
				Differentiating submucosal glial (70)	BCAN, APOE, LAMP5	No locational preference	Increases over time in colon and overall
				Glial progenitors (71)	PHOX2B, HMGA2, CTGF	No locational preference	Decreases over time in colon, TI and overall
		Neuronal	ELAVL4, CHRNA3, GAP43	ENS progenitors(72)	DLL3, DLL1, ELAVL4(low)	No locational preference	Decreases over time in colon and overall
				Neuroendocrine 1 (73)	NEUROD6, SCGN, SSTR2, NPY2R	No locational preference	Increases in time in colon and overall
				Neuroendocrine 2 (74)	DGKG, SST, KCTD16	No locational preference	Increases in time in colon, TI and overall
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1		İ	I			No locational preference	No significant differences
				Excitory motor neuron (75)	CASZ1, RAMP1, GFRA2	No locational preference	No significant differences
				Inhibitory motor neuron precursor (76)	SCGN, CRABP1, NOS1, VIP, ADCYAP1	No locational preference	Decreases over time in TI and overall
				Inhibitory motor neuron (77)	NOS1, VIP, HTR2B, ETV1	No locational preference	Increases then decreases in time in colon and overall
				Interneuron (78)	PENK, ACHE, TAC1, ONECUT2	No locational preference	Decreases over time
Immune	PTPRC	Macrophage	MERTK, CTSC, CTSD, CD14 (hi)	Macrophage (79)	FGL2, HES1, SLC40A1	Increased in colon (*)	Increases over time in colon and overall
				Macrophage SPP1+ (80)	SPP1, RNASE1, FOLR2	Increased in colon (*)	No significant differences
		Monocyte	CD14 (lo)	Monocyte (81)	FCN1, S100A8, CLEC12A	No locational preference	No significant differences
		Dendritic cell	ETV6, FLT3	pDCs (82)	IL3RA, LILRA4, PTCRA	No locational preference	No significant differences
				DCs (83)	CD207, BATF3, CD1D	No locational preference	No significant differences
		Adaptive lymphoid cells	CD19, EBF1, PAX5	Pre B Cell (84)	IGLL1, VPREB1, RAG1	No locational preference	No significant differences
				B cell (85)	BANK1, CD79B, IGHD	No locational preference	Increases over time in colon, TI and overall
		Innate lymphoid cells	ETS1, TOX	Naïve T Cell (86)	CD3D, CD3G, CD27	No locational preference	Increases over time in colon, TI and overall
				NKs and γδ (87)	CD8A, IL2RB, TRDC	Increased in small intestine (*)	No significant differences
				Type 3 ILCs (88)	IL7R, ID2, RORC, KRT86	No locational preference	Increases over time in colon and overall
		Eosinophils	N/A	Mast (89)	TPSAB1, CPA3, MS4A2	No locational preference	Increases over time in TI and overall. Decreases in colon.
		Cycling	N/A	Immune Cell cycle (90)	MKI67, CENPF, TOP2A	No locational preference	Decreases over time in colon, TI and overall
	MYH11, ACTG2, TAGLN	Cycling	N/A	Muscularis G2M phase (91)	MKI67, CENPF, TOP2A	No locational preference	Decreases then increases over time in colon
				Muscularis S-phase (92)	PCLAF, TYMS, GINS2	No locational preference	No significant differences
		Interstitial Cells	KIT	Interstitial Cells of Cajal / ICCs (93)	ANO1, ETV1, SPON2	No locational preference	No significant differences
				PDGFRA+ cells (94)	PDGFRA, ITGA8, BMP4	No locational preference	Decreases over time in colon and overall
_		Outer muscularis	FOXF2(-)	OM (95)	BCHE, CAPN3, IGFBP5	No locational preference	Increases over time in colon and overall
Muscle		Inner muscularis (IM)	FOXF2, ACTC1	Proximal IM (96)	IGFBP2, HOXC8, IGFBP7	Increased in small intestine (****)	No significant differences
cle				Distal IM (97)	PNCK, MORN5, AHNAK2	Increased in colon (***)	Increases over time in colon and overall
				Proximal MM (98)	PLCG2, NPNT, IGF1	Increased in small intestine (****)	No significant differences
				Distal MM (99)	PLN, MOXD1, TMEM158	Increased in colon (***)	Increases over time in colon and overall
				IM <i>PMAIP1</i> + (100)	PMAIP1, HMGCS1, KRT17	No locational preference	Increases then decreases over time in colon and overall. Increases in TI.
		IM progenitor	N/A	IM progenitor (101)	HOXC5, PLPP2, MYOCD	No locational preference	Decreases over time overall.