

## Online Resource 7: Summary of most significant networks

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*Lactobacillus paracasei* and *Lactobacillus plantarum* Strains Downregulate Proinflammatory Genes in an *Ex Vivo* System of Cultured Human Colonic Mucosa

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### **Authors:**

Christine Bäuerl<sup>a</sup>, Marta Llopis<sup>b</sup>, María Antolín<sup>b</sup>, Vicente Monedero<sup>a</sup>, Manuel Mata<sup>c</sup>, Manuel Zúñiga<sup>a</sup>, Francisco Guarner<sup>b</sup> and Gaspar Pérez Martínez<sup>a</sup>

### **Affiliation:**

<sup>a</sup> Department of Food Biotechnology, Instituto de Agroquímica y Tecnología de Alimentos, CSIC, Valencia, Spain

<sup>b</sup> Digestive System Research Unit, University Hospital Vall d'Hebron, Centro de Investigación Biomédica en Red de Enfermedades Hepáticas y Digestivas (Ciberehd), Barcelona, Spain

<sup>c</sup> Research Foundation, Hospital General Universitario, University of Valencia, Valencia, Spain

### **E-Mail of the corresponding Author Address:**

Gaspar Perez Martinez, e-mail: [gaspar.perez@iata.csic.es](mailto:gaspar.perez@iata.csic.es)

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**Table a:** Significant networks (Score≥ 15) derived from the up - and downregulated genes in Inflamed compared to Control sample.

Net-work	Molecules in network	Score	Focus molecules	Top functions
1	Ap1, <b>ATP2A3</b> , CCR3, CHEMOKINE; <b>CITA</b> , Creb, <b>CXCL5</b> , <b>CXCL6</b> , <b>CXCL9</b> , <b>CXCL11</b> , CXCR3, <b>EHMT2</b> , <b>ETS1</b> , <b>GABBR1</b> , <b>GBP2</b> , <b>GZMB</b> , Histone h3, Histone h4, <b>IFNG</b> , <b>IL2</b> , <b>IL17A</b> , <b>IL17F</b> , <b>IL2RA</b> , <b>IRF1</b> , <b>MCM4</b> , <b>NF1</b> , Nfat (family), NFκB (complex), <b>PIAS2</b> , REL/RELA/RELB, RNA polymerase II, <b>SERPINB2</b> , STAT5a/b, <b>TNFRSF4</b> , TNFSF4	39	21	Cell-to-Cell Signaling and Interaction, Inflammatory Response, Hematological System Development and Function
2	CXCL10, <b>CYB561</b> , DGKA, <b>FGF2</b> , FGFR1, <b>GBP1</b> , <b>GBP4</b> , heparin, <b>IDO1</b> , IFIT3, IFNAR2, <b>IL2</b> , <b>IL11</b> , IL15, <b>IL2RA</b> , IRF7, <b>IRF9</b> , LCK, MAGEA11, MIR31, <b>MX2</b> , <b>NUMB</b> , PF4, SLPI, <b>SPRYD5</b> , STAT1, <b>TNFRSF9</b> , TNFSF9, TP53, VAV1, <b>XCL1</b> , <b>XCL2</b> , XCR1, ZNF148, <b>ZNF655</b>	26	15	Cellular Growth and Proliferation, Cell-to-Cell Signaling and Interaction, Inflammatory Response,
3	<b>B3GAT1</b> , <b>C13ORF15</b> , <b>C1ORF144</b> , CADM1, <b>CD226</b> , CDC2, <b>CRTAM</b> , <b>EGR2</b> , ERBB2, FLOT2, FOSL1, FYN, GADD45, <b>GADD45G</b> , GBX2, GJB1, GLRA1, HMGA2, <b>HOPX</b> , ID4, MAPKBIP1, MIR9-1, <b>MR1</b> , <b>PIK3CB</b> , Plexin A, Plexin B, PLXNA1, PRX, <b>PTPN13</b> , REST, <b>RND1</b> , SLC18A3, SRC, SRF, TXK	21	12	Cell-to-Cell Signaling and Interaction, Neurological Disease, Cellular Development
4	A1CF, <b>ABCG8</b> , ANPEP, ANTXR1, APOB, ARFGAP1, CD55, CEBPB, COPB1, <b>CPN1</b> , <b>DMC1</b> , <b>FYCO1</b> , GBE1, GJB1, HNF1A, HNF4A, JUNB, LRP5, <b>LRP6</b> , <b>MGEA5</b> , NUF2, PEX13, PLG, PPP1CA, PPP1R2, PPP1R8, <b>PPP1R11</b> , PPP1R2P9, <b>PPP1R9A</b> , PPP2R5B, <b>PRG2</b> , SCD, SLPI, <b>STK19</b> , <b>WBP1</b>	18	11	Cardiovascular Disease, Lipid Metabolism, Molecular Transport

**Table b:** Significant networks (Score > 15) derived from the up - and downregulated genes in iLP compared to inflamed sample.

Net-work	Molecules in network	Score	Focus molecules	Top functions
1	Ap1, <b>ATP2A3, BAT2L</b> , Calbindin, CCR3, <b>CD59</b> , CHEMOKINE, Creb, <b>CXCL5, CXCL6, CXCL9, CXCL11, EHMT2, ELP3, ETS1, GZMB</b> , H3F3B, Histone h3, Histone h4, HIVEP3, <b>IFNG, IL2, IL17A, MCM4, NF1</b> , Nfat (family), NFκB (complex), REL/RELA/RELB, RNA polymerase II, <b>SERPINB2</b> , SRGN, TFIIA, THAP7, <b>TNFRSF4</b> , TNFSF4	31	18	Cell-to-Cell Signaling and Interaction, Hematological System Development and Function, Immune Cell Trafficking
2	<b>ABCG8</b> , ACTR5, ACTR8, ADSS, <b>CD226</b> , DHX8, DSN1, ERVK6, GATS, <b>GBE1</b> , HLA-B, HNF4A, INO80, INO80B, <b>INO80D</b> , INO80E, ITGAL, ITGB1, <b>LGALS8, LILRB3, MIS12</b> , NSL1, NUF2, PGD, PPARA, <b>PPP1R11, PRIC285, PTPN13, RPRD1B, RPS14</b> , RUVBL1, <b>SLC16A6, STK19, TPP2, ZNF317</b>	27	15	Endocrine System Disorders, Immunological Disease, Metabolic Disease
3	<b>B3GAT1</b> , CA11, COL11A1, <b>EHMT2, FGF2, FGF5, FGF9, FGF19, FGFR1, GATA1, GBP1, GBP4</b> , HDAC2, heparin, <b>HOPX, IDO1</b> , IFIT3, IFITM1, <b>INHBB</b> , Interferon Regulatory Factor, IRF7, <b>IRF9, LRDD, MX2, PRKAA2, RCOR1, REST, RIPK1, STMN2, TP53, TRIO, XCL1, XCL2</b> , XCR1, ZBP1	24	14	Gene Expression, Cellular Development, Cell Cycle
4	AHNAK, ALDOC, BAG1, <b>C1ORF144</b> , CSH1, DFFA, FOS, GRIN1, HIF1A, HTT, JUN/JUNB/JUND, <b>KDM3A, L3MBTL</b> , LCK, <b>LOX, LYST</b> , MIR17, MIRLET7A1, <b>PHF8, PHF13, PRRX1, RAI14</b> , RB1, RBP1, RTN1, <b>SEMA4B, SH3BP4, SLC4A7, SRGAP1, TNFRSF9</b> , UCP3, VSNL1, YWHAB, YWHAZ, <b>ZNF655</b>	23	14	Cell Morphology, Hematological System Development and Function, Cell Cycle

**Table c:** Significant networks (Score $\geq 15$ ) derived from the up - and downregulated genes in iLP(A $^{-}$ ) compared to inflamed sample.

Net-work	Molecules in network	Score	Focus molecules	Top functions
1	Ap1, <b>BAT2L, BCL10</b> , Caspase, CHEMOKINE, Creb, <b>CXCL5, CXCL6, CXCL9, CXCL11, EHMT2, ETS1, GZMB</b> , Histone h3, Histone h4, <b>HOXA9, IFNG, IL2, IL17A, IL2RA, IL7R, MCM4, MED6, MS12</b> , Nfat (family), NF $\kappa$ B (complex), REL/RELA/RELB, <b>RIPK1</b> , RNA polymerase II, <b>SERPINB2, STAT4, STAT5a/b, TNFRSF4</b> , TNSF4, WHSC1L1	41	22	Cellular Development, Cellular Growth and Proliferation, Inflammatory Response
2	A1CF, ANPEP, ARFGAP1, C12ORF11, CD55, <b>CD226, COPB1, CPN1, DUSP15</b> , EFS, ERRFI1, <b>FYCO1, FYN, GBE1, GJB1, GRB2, HIST2H2AA3, HNF1A, HNF4A, IL11, IL11RA, IL2RB, IL6ST, NUF2, ORC1L, PEX13, RAI14, SF3B4, SLC16A6, SLPI, STK19, VAV1, WBP1, ZNF317, ZNF655</b>	16	10	Cell Death, Cell-to-Cell Signaling and Interaction, Cell Morphology
3	APP, BACE1, BLMH, chondroitin sulfate C, Ck2, COL11A1, <b>FAM57A, FGF2, FGF5, FGF7, FGF9, FGF10, FGF19, FGFR4, FGFR1, FMR1, ganglioside GM1, GZMB, HABP2, heparan sulfate, heparin, LRP, MIR1-2, MIR29A, MIRLET7A1, NUFIP2, PAPD5, PAWR, PF4, PHF8, SLC4A7, TNRC6C, XCL1, XCL2, XCR1</b>	16	10	Cellular Assembly and Organization, Carbohydrate Metabolism, Small Molecule Biochemistry

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**Table d:** Significant networks (Score  $\geq 15$ ) derived from the up - and downregulated genes in iBL23 compared to inflamed sample.

Net-work	Molecules in network	Score	Focus molecules	Top functions
1	Ap1, <b>ATP2A3, BAT2L</b> , CCR3, CHEMOKINE, Creb, <b>CXCL5, CXCL6, CXCL9, CXCL11</b> , CXCR3, <b>EHMT2, ETS1, FSCN1, GABBR1, GZMB</b> , Histone h3, Histone h4, HIVEP3, <b>HOXA9, IFNG, IL2, IL12 (family), IL17A, MCM4, NF1</b> , Nfat (family), NF $\kappa$ B (complex), REL/RELA/RELB, RNA polymerase II, SRGN, <b>STAT4, TNFRSF4</b> , TNFSF4, WHSC1L1	32	18	Cell-to-Cell Signaling and Interaction, Inflammatory Response, Hematological System Development and Function
2	<b>CCNG2, CDK4, DGKA, EGR2, FN1, GATA1, GBP1, GBP4, heparin, ID2, IDO1, IFITM1, INHBB, Interferon Regulatory Factor, IRF7, IRF9, LCK, LOX, MR1, MX2, PITPNC1, PPT2, PRKAA2, PRRX1, RB1, TNFRSF9, TNFSF9, TP53, TRIO, XCL1, XCL2, XCR1, ZFB36, ZNF148, ZNF655</b>	30	17	Cell Cycle, Cellular Development, Cellular Growth and Proliferation
3	ARFGAP1, <b>B3GAT1, CD55, CD226, COPB1, DLG4, DSN1, FYN, GABBR1, GIPC2, HNF4A, IER5L, IL11, IL11RA, IL2RB, IL6ST, LPIN1, MIS12, NDC80, NSL1, NUF2, POU5F1, PPARA, PRIC285, REST, SCD, SEMA4B, SF3B4, SLC16A6, ST13, TCF7, WBP1, WIPF1, WWP2, ZNF317</b>	19	11	Lipid Metabolism, Molecular Transport, Small Molecule Biochemistry
4	ACTR5, <b>AIF1L, BLMH, BMP1, C1ORF144, CASP2, CASP3, CCDC50, CPA5, CTRL, CTSS, ERVK6, FGL2, GATS, HABP2, INO80, INO80D, INO80E, KDM3A, LRP6, MAP3K7IP3, MIR17, PAPPA, peptidase, PHF13, PPP1R11, PPP2R5C, RAI14, RIPK1, RIPK3, RNF130, RUVBL1, TAX1BP1, UBC, ZBP1</b>	17	10	Protein Degradation, Protein Synthesis, Reproductive System Disease