

Table 3. List of hypomethylated and upregulated genes and their reported associations with colorectal cancer (CRC) or other cancer types.

Gene	role	Linked to CRC or other cancers	Reference
Slc39a14	This gene encodes a member of the SLC39A family of divalent metal transporters that mediates the cellular uptake of manganese, zinc, iron, and cadmium.	The SLC39A14-exon4B transcript variant is a CRC biomarker with high sensitivity and organ-confined specificity.	DOI: 10.1002/ijc.27399
Map3k6	<i>MAP3K6</i> (Mitogen-Activated Protein Kinase Kinase Kinase 6) is a gene that encodes a kinase enzyme involved in signaling pathways that regulate various cellular processes, including cell growth, survival, and response to stress.	Germline mutations in <i>MAP3K6</i> are linked to familial gastric cancer	doi: <a href="https://doi.org/10.1371/journal.pgen.1004669">10.1371/journal.pgen.1004669</a>
Slc4a4	<i>SLC4A4</i> (Solute Carrier Family 4 Member 4) encodes a sodium bicarbonate cotransporter, which plays an important role in regulating pH balance within cells and across tissues, particularly in the kidneys and other epithelial tissues.	<i>SLC4A4</i> may serve as a novel biomarker for predicting prognosis in patients with colon adenocarcinoma.	doi: <a href="https://doi.org/10.12659/MSM.925016">10.12659/MSM.925016</a>
Arntl	Arntl is a core component of the circadian clock machinery and plays a key role in regulating various physiological processes, including metabolic and immune functions.	<i>BMAL1</i> (Arntl) is linked to bevacizumab resistance in colorectal cancer through the regulation of vascular endothelial growth factor A (VEGFA)	DOI: <a href="https://doi.org/10.1016/j.ebiom.2019.07.004">10.1016/j.ebiom.2019.07.004</a>
Myo6	<i>MYO6</i> (Myosin VI) is a motor protein that moves along actin filaments and plays critical roles in cellular processes such as endocytosis, secretion, and cell migration	<i>Overexpression of myosin VI promotes the progression of gastric cancer cells.</i>	<a href="https://doi.org/10.1016/j.gene.2016.08.015">https://doi.org/10.1016/j.gene.2016.08.015</a>
Matk	<i>MATK</i> (Megakaryocyte-Associated Tyrosine Kinase), also known as <i>CHK</i> , is a member of the C-terminal Src kinase (CSK) family.	<i>CSK-homologous kinase is a potential tumor suppressor gene in colorectal cancer, often epigenetically silenced through promoter methylation.</i>	<a href="https://doi.org/10.1038/s41388-021-01755-z">https://doi.org/10.1038/s41388-021-01755-z</a>

Pcyt1a	<i>PCYT1A (Phosphate Cytidylyltransferase 1, Choline, Alpha) is a gene encoding an enzyme involved in the synthesis of phosphatidylcholine, a major phospholipid in cell membranes.</i>	<i>PCYT1A suppresses proliferation and migration via inhibiting mTORC1 pathway in lung adenocarcinoma</i>	doi: 10.1016/j.bbrc.2020.05.164.
Cacna2d2	<i>CACNA2D2 (Calcium Voltage-Gated Channel Auxiliary Subunit Alpha2delta 2) is a gene that encodes a subunit of voltage-gated calcium channels, which play a crucial role in calcium influx regulation.</i>	<i>CACNA2D2 is a tumor suppressor gene in several types of cancer including CRC.</i>	doi: <a href="https://doi.org/10.18632/oncotarget.15815">10.18632/oncotarget.15815</a>
Bcas1	<i>BCAS1 is a candidate oncogene, amplified in various human cancers, including breast and prostate cancer, leading to more aggressive tumors.</i>	<i>BCAS1 inhibits <math>\beta</math>-Arrestin 2, thereby promoting the proliferation and migration of glioblastoma cells.</i>	<a href="https://doi.org/10.3390/cancers14163890">https://doi.org/10.3390/cancers14163890</a>
Dll1	<i>DLL1 (Delta-Like Ligand 1) is a ligand in the Notch signaling pathway, which is crucial for cell communication and regulates cell differentiation, proliferation, and apoptosis.</i>	<i>The Notch ligand DLL1 exhibits carcinogenic properties in human breast cancer cells.</i>	doi: <a href="https://doi.org/10.1371/journal.pone.0217002">10.1371/journal.pone.0217002</a>
Rxra	<i>RXRA is a nuclear receptor regulates gene expression by binding to retinoic acid and other ligands. This gene controls processes related to cell growth, differentiation, apoptosis, and metabolism.</i>	<i>RXRA influences the development of metachronous colorectal neoplasia and may be particularly important in the formation of proximal lesions.</i>	DOI: <a href="https://doi.org/10.1158/0008-5472.CAN-09-3264">10.1158/0008-5472.CAN-09-3264</a>
Ada	<i>Ada is an adenosine deaminase, which is essential for breaking down adenosine and deoxyadenosine in purine metabolism.</i>	<i>Polymorphic sites in ADA may impact cell-mediated anti-tumor immune responses by regulating adenosine levels and affecting additional protein functions beyond its enzymatic activity.</i>	<a href="https://doi.org/10.3109/07357907.2010.483501">https://doi.org/10.3109/07357907.2010.483501</a>
Hnf4a	<i>HNF4A is a nuclear transcription factor involved in regulating genes associated with metabolism, cell differentiation, and epithelial integrity.</i>	<i>In colorectal cancer, the transcription factor HNF4 has been associated with metastasis at both the RNA and protein levels.</i>	<a href="https://doi.org/10.1038/s41598-018-36168-8">https://doi.org/10.1038/s41598-018-36168-8</a>
Dpep1	<i>DPEP1 (Dipeptidase 1) is an enzyme primarily involved in the metabolism of dipeptides, particularly in the kidney, where it plays a role in breaking down certain peptides.</i>	<i>Dpep1 enhances the proliferation of colon cancer cells through regulation of the DPEP1/MYC feedback loop.</i>	doi: 10.1016/j.bbrc.2020.08.063.

Hkdc1	<i>HKDC1 (Hexokinase Domain Containing 1) is a member of the hexokinase family, enzymes that play an essential role in glycolysis by catalyzing the phosphorylation of glucose to glucose-6-phosphate.</i>	<i>Upregulation of HKDC1 promotes glycolysis and disease progression in gastric cancer, while also conferring chemoresistance.</i>	doi: <a href="https://doi.org/10.1111/cas.15692">10.1111/cas.15692</a>
Lrig3	<i>LRIG3 is part of the LRIG family, known for regulating receptor tyrosine kinase signaling pathways, including the EGF receptor pathway.</i>	<i>LRIG3 may play a role in regulating EGF receptor signaling and act as a tumor suppressor gene in glioma pathogenesis.</i>	doi: <a href="https://doi.org/10.3389/fonc.2019.00447">10.3389/fonc.2019.00447</a>
Ace	<i>ACE is a key enzyme in the renin-angiotensin system, primarily responsible for regulating blood pressure and fluid balance. ACE also influences cancer biology by promoting cell proliferation, angiogenesis, and tumor growth.</i>	<i>Polymorphism in the angiotensin I-converting enzyme gene is associated with tumor size and patient survival in colorectal cancer.</i>	doi: <a href="https://doi.org/10.1593/neo.07418">10.1593/neo.07418</a>
Fbp2	<i>FBP2 (Fructose-1,6-Bisphosphatase 2) is an enzyme involved in gluconeogenesis, the metabolic pathway that generates glucose from non-carbohydrate sources</i>	<i>Fbp2 inhibits sarcoma progression by restricting mitochondrial biogenesis.</i>	DOI: <a href="https://doi.org/10.1016/j.cmet.2019.10.012">10.1016/j.cmet.2019.10.012</a>
Pitx1	<i>PITX1 (Paired Like Homeodomain 1) is a transcription factor that plays an important role in regulating gene expression during development, particularly in limb and organ formation.</i>	<i>PITX1 plays a dual role in colon cancer, acting as both a tumor suppressor and an oncogene.</i>	doi: <a href="https://doi.org/10.3389/fonc.2023.1253238">10.3389/fonc.2023.1253238</a>
<i>lqgap2</i>	<i>Q motif-containing GTPase-activating protein 2 (IQGAP2) is a multidomain scaffolding protein that plays a significant role in various cellular processes, including cytoskeleton regulation, cell adhesion, and signal transduction.</i>	<i>lqgap2 mRNA serves as an independent prognostic marker and is associated with immunosuppression in diffuse large B-cell lymphoma (DLBCL).</i>	<a href="https://doi.org/10.1186/s12885-021-08086-y">https://doi.org/10.1186/s12885-021-08086-y</a>
Sema5a	<i>SEMA5A (Semaphorin 5A) is a member of the semaphorin family of proteins, which are involved in various aspects of cellular signaling, particularly in the nervous system.</i>	<i>A combined expression signature of ULBP2 and SEMA5A serves as a prognostic and predictive biomarker for colon cancer.</i>	doi: <a href="https://doi.org/10.7150/jca.17872">10.7150/jca.17872</a>

SLC38A4	SLC38A4 (Solute Carrier Family 38 Member 4) is a member of the solute carrier (SLC) family of transport proteins, which are critical for the transport of various solutes across cell membranes.	<i>SLC38A4 can be used as a prognostic biomarker for colorectal cancer.</i>	DOI: 10.1016/j.heliyon.2024.e34464
Cdx1	CDX1 (Caudal Type Homeobox 1) is a critical transcription factor for maintaining the identity and function of intestinal epithelial cells.	<i>Cdx1 overexpression could increase malignancy in early stages of tumourigenesis.</i>	<a href="https://doi.org/10.1038/onc.2008.78">https://doi.org/10.1038/onc.2008.78</a>
<i>Aldh1</i>	ALDH1 (Aldehyde Dehydrogenase 1) is an enzyme that plays a crucial role in the metabolism of aldehydes, particularly in the conversion of aldehydes to their corresponding carboxylic acids.	<i>Aldh1 could be a potential diagnostic and therapeutic target for colon cancer stem cells (CSCs).</i>	<a href="https://doi.org/10.3389/fonc.2022.1026278">https://doi.org/10.3389/fonc.2022.1026278</a>
<i>Pfkfb4</i>	Pfkfb4 (6-Phosphofructo-2-kinase/fructose-2,6-bisphosphatase 4) is an enzyme that plays a significant role in regulating glucose metabolism and energy homeostasis.	<i>High PFKFB4 expression was associated with both improved overall survival (OS) and post-progression survival (PPS) in Colon adenocarcinoma patients.</i>	<a href="https://doi.org/10.1038/s41598-023-43619-4">https://doi.org/10.1038/s41598-023-43619-4</a>
<i>Mgat4a</i>	MGAT4A (Mannosyl (alpha-1,3-)-glycoprotein beta-1,4-N-acetylglucosaminyltransferase A) is an enzyme involved in glycoprotein biosynthesis.	<i>Aberrant expression of MGAT4A mRNA has been observed in various cancer cells and is known to enhance their invasiveness by modulating the functions of glycoproteins, including integrin <math>\beta</math>1.</i>	<a href="https://doi.org/10.1038/s42003-022-03661-w">https://doi.org/10.1038/s42003-022-03661-w</a>
<i>Aldh9a1</i>	ALDH9A1 (Aldehyde Dehydrogenase 9 Family Member A1) is an enzyme that plays a crucial role in the metabolism of aldehydes, particularly in the detoxification of harmful substances and the regulation of cellular processes.	<i>The abundance of ALDH9A1, is elevated in renal cell carcinoma (RCC).</i>	<a href="https://doi.org/10.1074/mcp.M800252-MCP200">https://doi.org/10.1074/mcp.M800252-MCP200</a>
<i>Chmp4c</i>	<i>Chmp4c</i> (Charged Multivesicular Body Protein 4C) is a member of the CHMP family of proteins, which are involved in the endosomal sorting complex required for transport (ESCRT) machinery.	<i>Chmp4c plays a role in the viability and motility of cervical cancer (CC) cells by regulating epithelial-mesenchymal transition, potentially aiding in the discovery of novel biomarkers for CC therapy.</i>	doi: <a href="https://doi.org/10.1002/2211-5463.12880">10.1002/2211-5463.12880</a>

<i>EPB41L1</i>	<i>EPB41L1 (Erythrocyte Membrane Protein Band 4.1 Like 1) is a member of the band 4.1 family of proteins, which are known for their roles in maintaining the integrity and stability of the cytoskeleton in various cell types.</i>	<i>EPB41L1 is significantly upregulated in colorectal adenocarcinoma tissues, and its high expression level is associated with a poor prognosis for the patients.</i>	DOI: <a href="https://doi.org/10.21203/rs.3.rs-358130/v1">https://doi.org/10.21203/rs.3.rs-358130/v1</a>
<i>SNX7</i>	<i>SNX7 (Sorting Nexin 7) is a member of the sorting nexin family of proteins, which are involved in intracellular trafficking and membrane dynamics.</i>	<i>SNX7 is emerging as a promising novel biomarker for the diagnosis, prognosis, and prediction of treatment responses to chemotherapy and immunotherapy in patients with hepatocellular carcinoma (HCC).</i>	doi: <a href="https://doi.org/10.1186/s12885-023-11405-0">10.1186/s12885-023-11405-0</a>
<i>Coro2a</i>	<i>CORO2A (Coronin 2A) is a protein that plays a role in the regulation of the actin cytoskeleton and is involved in various cellular processes.</i>	<i>CORO2A has the potential to serve as a prognostic predictor for patients with triple-negative breast cancer (TNBC).</i>	doi: <a href="https://doi.org/10.3389/fonc.2020.00916">10.3389/fonc.2020.00916</a>
<i>Dab1</i>	<i>DAB1 (Disabled 1) is a protein that plays a critical role in various cellular signaling pathways, particularly in neuronal development and migration.</i>	<i>Genetic depletion of DAB1 inhibits cancer invasion and metastasis in mice with activated NOTCH signaling.</i>	DOI: <a href="https://doi.org/10.1158/2159-8290.CD-14-0595">10.1158/2159-8290.CD-14-0595</a>
<i>Rbp7</i>	<i>RBP7 (RNA Binding Protein 7) is a protein involved in various cellular processes, particularly in RNA metabolism and gene regulation.</i>	<i>RBP7 is a robust prognostic biomarker in colon cancer and functionally contributes to the malignant characteristics of colon cancer cells.</i>	doi: <a href="https://doi.org/10.7150/jca.35180">10.7150/jca.35180</a>
<i>Ugdh</i>	<i>UGDH (UDP-glucuronic acid decarboxylase) is an enzyme involved in the metabolism of carbohydrates and the detoxification of various compounds in the body.</i>	<i>UGDH is an enzyme that could be utilized as a potential prognostic marker in oncology and as a therapeutic target in cancer biology.</i>	doi: <a href="https://doi.org/10.18632/oncotarget.28514">10.18632/oncotarget.28514</a>
<i>Tpk1</i>	<i>TPK1 (Thiamine Phosphate Kinase 1) is an enzyme involved in thiamine metabolism and plays a crucial role in the synthesis of thiamine diphosphate.</i>	<i>TPK1 expression is a significant factor influencing the susceptibility of cancer cells to ionizing radiation.</i>	<a href="https://doi.org/10.18632">https://doi.org/10.18632</a>
<i>Gfpt1</i>	<i>GFPT1 (Glutamine-Fructose-6-Phosphate Transaminase 1) is an enzyme that plays a key role in amino acid and carbohydrate metabolism.</i>	<i>The enhanced levels of Gfpt1 promotes the growth of colorectal cancer.</i>	doi: <a href="https://doi.org/10.3390/ijms25094883">10.3390/ijms25094883</a> .

<i>Nadsyn1</i>	NADSYN1 (NAD synthetase 1) is an enzyme that catalyzes the final step in the biosynthesis of nicotinamide adenine dinucleotide (NAD+)	The DNA copy number of Nadsyn1 was found to be elevated in several cancer types, including prostate, ovarian, and pancreatic cancers.	doi: <a href="https://doi.org/10.1038/s41586-019-1150-2">10.1038/s41586-019-1150-2</a>
<i>Lrrc1</i>	The LRRC1 gene has the potential to function as an immune-associated prognostic biomarker for hepatocellular carcinoma (HCC).	The LRRC1 gene has the potential to function as an immune-associated prognostic biomarker for HCC.	doi: <a href="https://doi.org/10.1097/MD.00000000000034365">10.1097/MD.00000000000034365</a>
<i>Car12</i>	CAR12 (Carbonic Anhydrase 12) is an enzyme that catalyzes the reversible hydration of carbon dioxide to bicarbonate and a proton, playing a crucial role in maintaining acid-base balance and facilitating various physiological processes.	Car12 is upregulated in human as well as murine breast cancer tissue.	<a href="https://doi.org/10.1186/s13058-023-01644-1">https://doi.org/10.1186/s13058-023-01644-1</a>
<i>Mst1r</i>	MST1R (Macrophage Stimulating 1 Receptor) or RON, is a receptor tyrosine kinase involved in various cellular processes.	RON plays a role in carcinogenesis through its immune modulation of the tumor microenvironment, activation of various oncogenic pathways, and protective functions under cellular stress.	doi: <a href="https://doi.org/10.3390/cancers14082037">10.3390/cancers14082037</a>
<i>Tspan8</i>	Tetraspanin 8 (TSPAN8) is a member of the tetraspanin family, characterized by four transmembrane domains and involvement in various cellular processes.	TSPAN8 enhances the stemness of cancer cells through the activation of sonic Hedgehog signaling.	<a href="https://doi.org/10.1038/s41467-019-10739-3">https://doi.org/10.1038/s41467-019-10739-3</a>
<i>Cellf5</i>	CELF (CUGBP Elav-like family) proteins are RBPs (RNA-binding proteins) with pleiotropic capabilities in RNA processing.	Members of the CELF family have been linked to widespread changes in cancer proliferation and invasion, resulting in their identification as potential tumor suppressors or even oncogenes.	doi: <a href="https://doi.org/10.3390/ijms222011056">10.3390/ijms222011056</a>
<i>Dpp4</i>	Dipeptidyl peptidase-4 (DPP-4), also known as CD26, is a multifunctional enzyme involved in various physiological processes.	DPP-4 inhibitors enhance the incidence of newly developed metastasis in primary thyroid cancer among diabetic patients.	doi: 10.1002/ijc.31870.

<i>Slc12a8</i>	SLC12A8 (Solute Carrier Family 12 Member 8) is a protein-coding gene that encodes a cation-chloride cotransporter.	The upregulation of SLC12A8 promoted resistance to apoptosis, EMT, and reduced reactive oxygen species (ROS) production.	doi: <a href="https://doi.org/10.21037/tcr-24-87">10.21037/tcr-24-87</a>
<i>Aldh1b1</i>	Aldehyde dehydrogenase 1B1 (ALDH1B1) is an enzyme that plays a crucial role in the metabolism of aldehydes, particularly in the detoxification of alcohol-derived acetaldehyde.	ALDH1B1 is significantly upregulated in colorectal adenocarcinoma, and silencing ALDH1B1 inhibits the growth of human colon cancer spheroids and xenografts.	<a href="https://doi.org/10.1371/journal.pone.0121648">https://doi.org/10.1371/journal.pone.0121648</a>
<i>Krt20</i>	Keratin 20 (KRT20) is a type I intermediate filament protein predominantly expressed in the gastrointestinal epithelium, including the colon.	Colorectal cancer cells typically exhibit high levels of keratin 20 (K20) expression	DOI: <a href="https://doi.org/10.14670/HH-27.347">10.14670/HH-27.347</a>
<i>Chst4</i>	Carbohydrate sulfotransferase 4 (CHST4) is an enzyme that catalyzes the transfer of sulfate groups to specific carbohydrate structures	CHST4 functions as a tumor suppressor in hepatocellular carcinoma associated with hepatitis B virus (HCC-HBV).	doi: <a href="https://doi.org/10.3389/fonc.2020.554331">10.3389/fonc.2020.554331</a>
<i>Sec24a</i>	SEC24A (SEC24 Homolog A) is a protein that plays a crucial role in the transport of proteins from the endoplasmic reticulum (ER) to the Golgi apparatus.	SEC24A is involved in enhancing the invasion and metastasis of human gastric cancer (GC) cells.	PMCID: PMC6962814 PMID: <a href="https://pubmed.ncbi.nlm.nih.gov/31949794/">31949794</a>
<i>Arhgap26</i>	ARHGAP26 (Rho GTPase-activating protein 26) is an enzyme that regulates the activity of Rho GTPases	The recurrent RhoGAP gene fusion CLDN18-ARHGAP26 enhances RHOA activation and stimulates focal adhesion kinase and YAP-TEAD signaling in diffuse gastric cancer.	<a href="https://doi.org/10.1136/gutjnl-2023-329686">https://doi.org/ 10.1136/gutjnl-2023-329686</a>
<i>Aff3</i>	AFF3 (AF4/FMR2 family member 3) is a gene encoding a nuclear transcriptional activator involved in RNA elongation and gene expression regulation.	AFF3 may be a promising potential marker for the diagnosis and prognosis of GC patients.	doi: <a href="https://doi.org/10.1002/jcla.24437">10.1002/jcla.24437</a>
<i>Rgs22</i>	Regulator of G-protein signaling 22 (RGS22) is a protein that modulates G-protein signaling pathways	RGS22 is overexpressed in low-metastatic epithelial cancers and plays a role in cell migration and invasion processes in esophageal cancer cell lines.	DOI: <a href="https://doi.org/10.1007/s10585-011-9390-z">10.1007/s10585-011-9390-z</a>

<i>Kctd12</i>	KCTD12 (Potassium Channel Tetramerization Domain Containing 12) is a protein-coding gene involved in various cellular processes, including the regulation of GABA-B receptor signaling and potential roles in cancer progression.	KCTD12 is a novel regulator of colorectal cancer (CRC) cell stemness and may act as a promising prognostic marker and therapeutic target for CRC patients.	<a href="https://doi.org/10.1038/srep20460">https://doi.org/10.1038/srep20460</a>
<i>Fyb2</i>	FYB2, also known as FYN binding protein 2, is an adaptor protein involved in T-cell receptor (TCR)-mediated activation of signaling pathways.	FYB2 may serve as a novel prognostic biomarker in liver hepatocellular carcinoma (LIHC)	<a href="https://doi.org/10.3390/livers2040027">https://doi.org/10.3390/livers2040027</a>
<i>Ighg3</i>	IGHG3 contributes to the structure of IgG3 antibodies, which are crucial for immune defense against pathogens.	Ighg3 is overexpressed in non-small cell lung carcinoma.	doi: 10.3892/ijo.26.1.247.
<i>Frat1</i>	FRAT1 (FRAT Regulator of WNT Signaling Pathway 1) is a protein-coding gene that plays a pivotal role in the Wnt signaling pathway, which is essential for regulating cell proliferation, differentiation, and survival.	Knocking out the FRAT1 gene reduces the expression of $\beta$ -catenin, cyclin D1, and c-myc in liver cancer cells, inhibiting tumor cell proliferation by modulating the Wnt/ $\beta$ -catenin signaling pathway.	doi: 10.3892/or.2016.5130.
<i>Rnase1</i>	RNase L is activated by 2-5A molecules, which are synthesized in response to viral infections.	RNase L is a crucial element in interferon (IFN)-mediated antiviral signaling and has demonstrated antitumor effects in cancer.	<a href="https://doi.org/10.1038/s41419-019-1902-9">https://doi.org/10.1038/s41419-019-1902-9</a>
<i>Syne2</i>	SYNE2 (Spectrin Repeat Containing Nuclear Envelope Protein 2), also known as Nesprin-2, is a protein that plays a crucial role in maintaining the structural integrity of the nucleus by anchoring it to the actin cytoskeleton.	SYNE2 plays a significant role in the regulation of p21 and is associated with the clinical outcomes of HBV-related hepatocellular carcinoma (HCC) in a TP53-independent manner.	<a href="https://doi.org/10.1038/srep31237">https://doi.org/10.1038/srep31237</a>
<i>Kalrn</i>	Kalirin (KALRN) is a gene encoding a protein that functions as a Rho guanine nucleotide exchange factor (RhoGEF), playing a pivotal role in regulating the actin cytoskeleton and influencing neuronal development and synaptic plasticity	The KALRN mutation serves as a valuable biomarker for predicting patients' responses to immunotherapy in cancer.	doi: <a href="https://doi.org/10.1136/jitc-2019-000293">10.1136/jitc-2019-000293</a>



<i>Nptx2</i>	Neuronal Pentraxin 2 (NPTX2) is a protein involved in synaptic function and plasticity within the central nervous system.	NPTX2 enhances the growth of colorectal cancer and liver metastasis through the activation of the canonical Wnt/ $\beta$ -catenin pathway via FZD6.	<a href="https://doi.org/10.1038/s41419-019-1467-7">https://doi.org/10.1038/s41419-019-1467-7</a>
<i>Itpk1</i>	Inositol-tetrakisphosphate 1-kinase (ITPK1) is an enzyme that plays a crucial role in inositol phosphate signaling pathways, which are essential for various cellular processes, including cell proliferation, differentiation, and apoptosis.	ITPK1 enhances the sensitivity of tumor cells to IgA-dependent neutrophil-mediated killing in vivo.	doi: 10.4049/jimmunol.2300372.
<i>Sh3rf2</i>	SH3RF2 (SH3 Domain Containing Ring Finger 2) is a protein-coding gene that encodes an E3 ubiquitin-protein ligase involved in various cellular processes, including apoptosis regulation and protein stability.	SH3RF2 acts as an oncogene by regulating the stability of the PAK4 protein.	doi: 10.1093/carcin/bgt338.
<i>Naaladl1</i>	NAALADL1 (N-acetylated alpha-linked acidic dipeptidase-like 1) is a protein-coding gene that encodes an aminopeptidase enzyme involved in peptide catabolism.	NAALADL1 gene expression is low in typical prostate cancer cell lines but is elevated in the human medullary thyroid carcinoma TT cell line.	doi: <a href="https://doi.org/10.1073/pnas.2025710119">10.1073/pnas.2025710119</a>
<i>Clrn3</i>	CLRN3 (Clarín 3) is a protein-coding gene located on chromosome 10, known for its involvement in lysosomal function and cellular processes.	CLRN3 significantly reduced the malignant biological behavior of colorectal cancer (CRC) cells	<a href="https://doi.org/10.3390/biom14080891">https://doi.org/10.3390/biom14080891</a>
<i>Pls1</i>	Plastin 1, also known as I-plastin or fimbrin, is an actin-bundling protein encoded by the PLS1 gene.	Plastin 1 promotes the metastasis of colorectal cancer by activating the IQGAP1/Rac1/ERK pathway.	doi: <a href="https://doi.org/10.1111/cas.14438">10.1111/cas.14438</a>
<i>Fcgbp</i>	FCGBP (Fc gamma binding protein) is a protein involved in mucosal immunity and has been implicated in various cancers.	FCGBP is a prognostic biomarker that is associated with immune infiltration in glioma.	<a href="https://doi.org/10.3389/fonc.2021.769033">https://doi.org/10.3389/fonc.2021.769033</a>
<i>Liph</i>	Lipase H (LIPH) is a membrane-bound enzyme that catalyzes the production of 2-acyl lysophosphatidic acid (LPA), a	LIPH (lipase H) is significantly overexpressed in breast cancer tissues and is associated with poorer prognoses in breast cancer.	doi: <a href="https://doi.org/10.2147/IJGM.S332233">10.2147/IJGM.S332233</a>

	bioactive lipid mediator involved in various physiological processes.		
<i>GPR133</i> ( <i>ADGRD1</i> )	GPR133, also known as ADGRD1, is an adhesion G protein-coupled receptor (aGPCR) involved in various cellular processes, including tumor progression.	GPR133 is a key mediator of the hypoxic response in glioblastoma (GBM) and plays significant protumorigenic roles.	<a href="https://doi.org/10.1038/oncsis.2016.63">https://doi.org/10.1038/oncsis.2016.63</a>
<i>Dcst1</i>	DCST1 (DC-STAMP Domain Containing 1) is a protein-coding gene that encodes a protein involved in various cellular processes, including immune response and cell signaling.	DCST1-AS1 enhances TGF- $\beta$ -induced epithelial-mesenchymal transition and increases chemoresistance in triple-negative breast cancer cells through the action of ANXA1.	doi: <a href="https://doi.org/10.3389/fonc.2020.00280">10.3389/fonc.2020.00280</a>
<i>Gucy2c</i>	Guanylate cyclase 2C (GUCY2C) functions as a receptor for endogenous peptides such as guanylin and uroguanylin, as well as for heat-stable enterotoxins produced by certain pathogens.	GUCY2C is a promising target for immunotherapy and serves as a diagnostic marker for both primary and metastatic disease.	doi: <a href="https://doi.org/10.1080/14728222.2021.1937124">10.1080/14728222.2021.1937124</a>
<i>Fcmr</i>	The FCMR gene encodes the Fc mu receptor, a protein that binds to the Fc region of immunoglobulin M (IgM) antibodies.	Therapeutically targeting Fcmr during oncogenesis reduced tumor growth, whether used alone or in combination with anti-PD-1.	<a href="https://doi.org/10.1038/s41467-019-10619-w">https://doi.org/10.1038/s41467-019-10619-w</a>
<i>Tnfrsf14</i>	TNFRSF14, also known as Herpesvirus Entry Mediator (HVEM), is a member of the tumor necrosis factor (TNF) receptor superfamily.	TNFRSF14 (HVEM) is a promising new immune checkpoint for cancer immunotherapy in humanized mice.	<a href="https://doi.org/10.1101/711119">https://doi.org/10.1101/711119</a>
<i>Dao</i>	Diamine oxidase (DAO), also known as amine oxidase, copper-containing 1 (AOC1), is an enzyme responsible for the oxidative deamination of biogenic amines, including histamine, putrescine, and spermidine.	Plasma diamine oxidase activity serves as a biomarker for assessing gastrointestinal tract toxicities in gastric cancer patients undergoing chemotherapy.	<a href="https://doi.org/10.1159/000336799">https://doi.org/10.1159/000336799</a>
<i>Sytl3</i>	Synaptotagmin-like 3 (SYTL3) acts as a peripheral membrane protein that interacts with RAB27A and is involved in the process of vesicular trafficking, facilitating the transport and release of cellular vesicles.	SYTL3 functions as a regulator of cortical neuronal migration in the developing brains of both humans and mice.	<a href="https://doi.org/10.1016/j.celrep.2021.108802">https://doi.org/10.1016/j.celrep.2021.108802</a>

<i>Ak7</i>	Adenylate kinase 7 (AK7) catalyzes the reversible phosphorylation of adenine nucleotides, playing a crucial role in maintaining cellular energy homeostasis.	Adenylate kinase 7 serves as a prognostic indicator for overall survival in ovarian cancer.	doi: <a href="https://doi.org/10.1097/MD.00000000000024134">10.1097/MD.00000000000024134</a>
<i>Bach2</i>	BACH2 (BTB Domain And CNC Homolog 2) is a transcription factor that plays a pivotal role in regulating immune responses and has been implicated in various cancers.	The transcription factor BACH2 facilitates tumor immunosuppression.	DOI: <a href="https://doi.org/10.1172/JCI82884">10.1172/JCI82884</a>
<i>Abcc8</i>	ABCC8, known as ATP-binding cassette sub-family C member 8, encodes the sulfonylurea receptor 1 (SUR1) protein, a subunit of the ATP-sensitive potassium (K-ATP) channel.	ABCC8 mRNA expression may serve as an independent prognostic indicator for glioma patients and could predict their sensitivity to temozolomide treatment.	doi: <a href="https://doi.org/10.1038/s41598-020-69676-7">10.1038/s41598-020-69676-7</a>
<i>Anpep</i>	ANPEP, also known as alanyl aminopeptidase, is a membrane-bound enzyme plays a crucial role in peptide digestion and metabolism.	ANPEP is a macrophage-related gene that may play a significant role in tumor metabolism.	<a href="https://doi.org/10.1158/1538-7755.DISP21-PO-153">https://doi.org/10.1158/1538-7755.DISP21-PO-153</a>
<i>Rap1gap2</i>	Rap1GAP2 (Rap1 GTPase-activating protein 2) functions as a GTPase-activating protein (GAP) for Rap1, a small GTPase involved in regulating cell adhesion, migration, and proliferation.	Rap1GAP may play a role in tumor progression in colorectal cancer (CRC) and could serve as a potential target for prognostic prediction in CRC patients.	<a href="https://doi.org/10.3892/ot.2018.8305">https://doi.org/10.3892/ot.2018.8305</a>
<i>Dgki</i>	Diacylglycerol kinase iota (DGKI) is part of the type IV diacylglycerol kinase subfamily which plays a crucial role in regulating intracellular levels of diacylglycerol (DAG) and phosphatidic acid (PA)	DGKI overexpression may serve as a potential molecular marker for poor prognosis in gastric cancer (GC), with the MAPK signaling pathway identified as a key pathway associated with DGKI regulation in this context.	<a href="https://doi.org/10.3389/fmed.2020.00320">https://doi.org/10.3389/fmed.2020.00320</a>
<i>Nos1ap</i>	Nitric oxide synthase 1 adaptor protein (NOS1AP), functions as an adaptor protein that interacts with neuronal nitric oxide synthase (nNOS), playing a crucial role in neuronal signaling and synaptic function.	A protein complex consisting of SCRIB, NOS1AP, and VANG1 regulates cell polarity and migration and is linked to the progression of breast cancer.	DOI: <a href="https://doi.org/10.1038/onc.2011.528">10.1038/onc.2011.528</a>
<i>Trpm4</i>	Transient Receptor Potential Melastatin 4 (TRPM4) is a calcium-activated, non-selective cation channel that allows the influx of monovalent cations, such as	TRPM4 is significantly expressed in human colorectal tumor buds and plays a role in the proliferation, cell cycle	<a href="https://doi.org/10.1002/1878-0261.12566">https://doi.org/10.1002/1878-0261.12566</a>

	sodium and potassium, while being impermeable to calcium.	regulation, and invasion of colorectal cancer cells.	
<i>Slc43a2</i>	SLC43A2, is a protein that functions as a transporter for large neutral amino acids, including methionine. It plays a crucial role in cellular amino acid uptake, influencing various metabolic processes.	SLC43A2 has been identified as an oncogene that regulates methionine metabolism and tumor proliferation across various cancer types.	doi: 10.1038/s41586-020-2682-1.
<i>Atp6v0a2</i>	ATP6V0A2, is a component of the vacuolar-type H <sup>+</sup> -ATPase (V-ATPase) complex, which is responsible for acidifying intracellular compartments such as endosomes, lysosomes, and the Golgi apparatus.	The ATP6V0A2 subunit is overexpressed in ovarian cancer and is linked to cell membrane dynamics that drive metastasis.	<a href="https://doi.org/10.1038/s41419-021-04495-w">https://doi.org/10.1038/s41419-021-04495-w</a>
<i>A4gnt</i>	Alpha-1,4-N-acetylglucosaminyltransferase (A4GNT) is an enzyme responsible for adding N-acetylglucosamine (GlcNAc) to glycoproteins, a modification that influences protein function and stability.	A4gnt knockout mice, the absence of αGlcNAc in gland mucin leads to the development of gastric adenocarcinoma.	doi: <a href="https://doi.org/10.1111/cas.12305">10.1111/cas.12305</a>
<i>Crybg2</i>	Crystallin beta-gamma domain-containing protein 2 is predicted to possess carbohydrate binding activity.	<i>The association of Crybg2 with colon cancer has not yet been elucidated.</i>	<a href="https://www.ncbi.nlm.nih.gov/gene/55057">https://www.ncbi.nlm.nih.gov/gene/55057</a>
<i>Slc5a4b</i>	Predicted to facilitate low-affinity D-glucose symporter activity, function upstream of or within proton transmembrane transport.	<i>The association of Slc5a4b with colon cancer has not yet been elucidated.</i>	<a href="https://www.ncbi.nlm.nih.gov/gene/6527">https://www.ncbi.nlm.nih.gov/gene/6527</a>
<i>Slc4a8</i>	t <i>Slc4a8</i> transports sodium and bicarbonate ions across cell membranes	<i>The association of Slc4a8 with colon cancer has not yet been elucidated.</i>	<a href="https://doi.org/10.1016/j.mam.2012.07.009">https://doi.org/10.1016/j.mam.2012.07.009</a>
<i>Myo7b</i>	Myo7b, or myosin VIIb, is a protein found in the brush border microvilli of epithelial cells in the colon and kidneys	<i>The association of Myo7b with colon cancer has not yet been elucidated.</i>	doi: <a href="https://doi.org/10.1016/j.cub.2016.08.014">10.1016/j.cub.2016.08.014</a>

<i>Epb41l4b</i>	PB41L4B (erythrocyte membrane protein band 4.1 like 4B) is a gene that encodes a protein involved in regulating the circumferential actomyosin belt in epithelial cells.	<i>Epb41l4b</i> is downregulated in CRC.	doi: <a href="https://doi.org/10.1042/BSR20200141">10.1042/BSR20200141</a>
<i>Espn</i>	Espin (encoded by the ESPN gene) is a multifunctional actin-bundling protein crucial for the organization and function of actin filament-rich structures, such as microvilli in mechanosensory and chemosensory cells.	High expression levels of ESPN are independently linked to a poor prognosis in patients with esophageal squamous cell carcinoma (ESCC), while the downregulation of ESPN inhibits the growth of ESCC cells.	<a href="https://doi.org/10.1186/s12935-018-0713-x">https://doi.org/10.1186/s12935-018-0713-x</a>
<i>Shroom3</i>	SHROOM3, a gene linked to chronic kidney disease, influences the structure of podocytes.	<i>The association of Shroom3 with colon cancer has not yet been elucidated.</i>	<a href="https://doi.org/10.1038/s41598-020-77952-9">https://doi.org/10.1038/s41598-020-77952-9</a>
<i>Atp2b2</i>	ATP2B2, also known as plasma membrane calcium-transporting ATPase 2 (PMCA2), is a gene encoding a protein responsible for regulating calcium ion concentrations within cells.	<i>The association of Atp2b2 with colon cancer has not yet been elucidated.</i>	doi: <a href="https://doi.org/10.1007/s00439-018-1965-1">10.1007/s00439-018-1965-1</a>
<i>2200002D01Rik</i>	The gene 2200002D01Rik is a protein-coding gene in mice, also known as RIKEN cDNA 2200002D01 gene.	<i>The association of 2200002D01Rik with colon cancer has not yet been elucidated.</i>	<a href="https://www.ncbi.nlm.nih.gov/gene/72275">https://www.ncbi.nlm.nih.gov/gene/72275</a>
<i>Dnajc13</i>	DNAJC13 is an endosome-related protein and believed to regulate endosomal membrane trafficking.	<i>The association of Dnajc13 with colon cancer has not yet been elucidated.</i>	<a href="https://doi.org/10.1093/hmg/ddy003">https://doi.org/10.1093/hmg/ddy003</a>
<i>Leap2</i>	Liver-expressed antimicrobial peptide 2 (LEAP2) is a cysteine-rich, cationic antimicrobial peptide predominantly expressed in the liver.	<i>The association of Leap2 with colon cancer has not yet been elucidated.</i>	<a href="https://doi.org/10.1016/j.fsi.2023.109225">https://doi.org/10.1016/j.fsi.2023.109225</a>
<i>Fsd1l</i>	FSD1L (Fibronectin Type III and SPRY Domain Containing 1 Like) is a protein-coding gene associated with hydrocephalus, a condition characterized by the accumulation of cerebrospinal fluid in the brain.	<i>The association of Fsd1l with colon cancer has not yet been elucidated.</i>	<a href="https://www.genecards.org/cgi-bin/carddisp.pl?gene=FSD1L">https://www.genecards.org/cgi-bin/carddisp.pl?gene=FSD1L</a>

<i>Itgad</i>	The ITGAD gene encodes the integrin alpha D protein, a component of the beta-2 integrin family of membrane glycoproteins.	<i>The association of Itgad with colon cancer has not yet been elucidated.</i>	<a href="https://www.genecards.org/cgi-bin/carddisp.pl?gene=ITGAD">https://www.genecards.org/cgi-bin/carddisp.pl?gene=ITGAD</a>
<i>Gm36640</i>	Gm36640 is a predicted gene located on mouse chromosome 6, spanning positions 135,256,550 to 135,290,997 on the positive strand. It is classified as a long non-coding RNA (lncRNA) gene.	<i>The association of Gm36640 with colon cancer has not yet been elucidated.</i>	<a href="https://www.informatics.jax.org/marker/MGI%3A5595799">https://www.informatics.jax.org/marker/MGI%3A5595799</a>