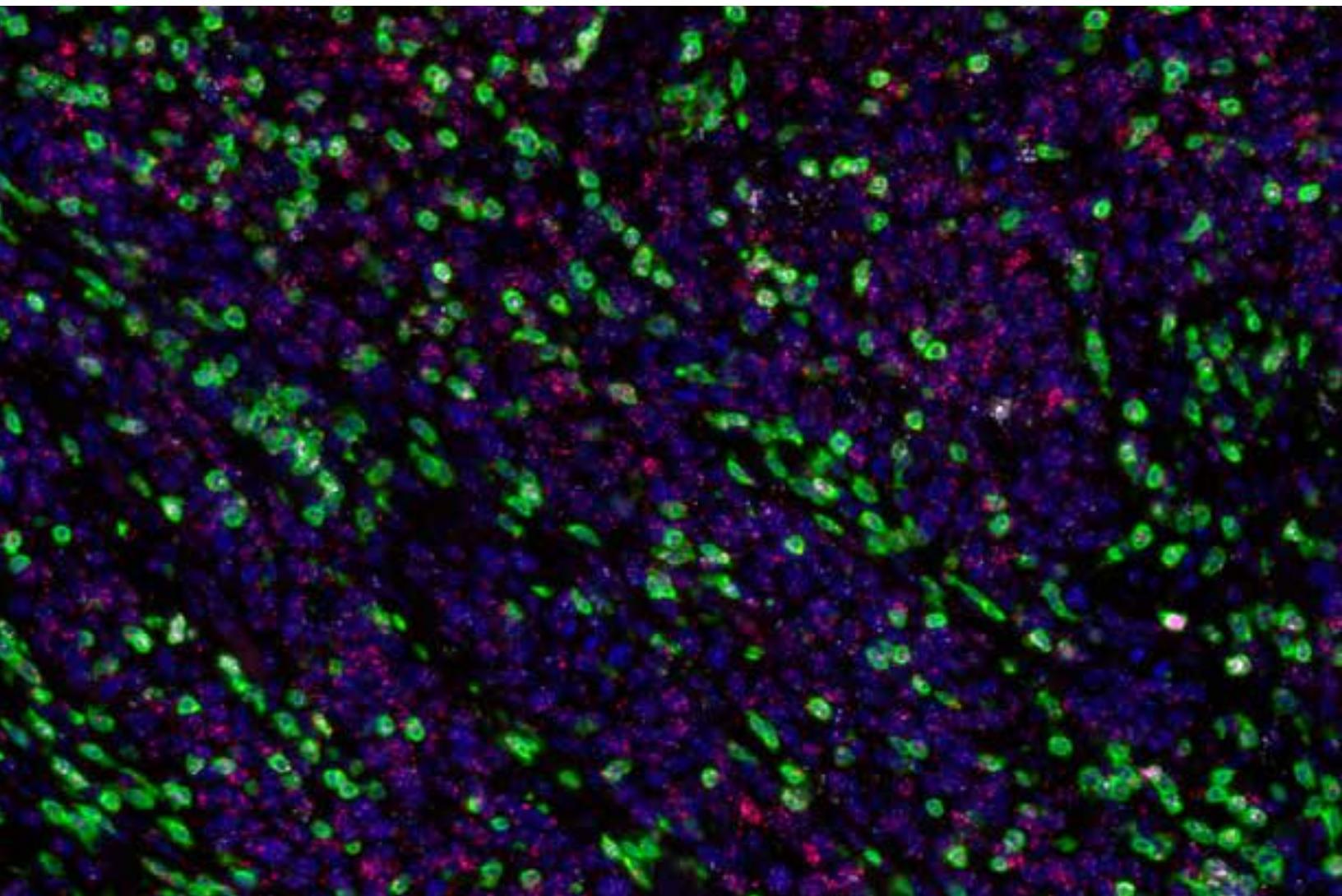


Learn about the Antibodies Researchers are Using with the RNAscope™ Technology

Publications utilizing the RNAscope *in situ*
hybridization technology in combination with
immunohistochemistry or immunofluorescence



Antigens

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Introduction

Spatially resolved gene expression has emerged as a crucial technique to understand complex multicellular interactions within the tissue. Traditional methods such as RNA *in situ* hybridization (ISH) and immunohistochemistry (IHC) or immunofluorescence (IF) allow gene expression analysis with spatial resolution at the RNA and protein level, respectively. Traditional RNA ISH has been challenging due to low sensitivity and specificity, as well as time-consuming and cumbersome to perform. The highly sensitive and specific RNAscope™ technology has been developed to overcome the limitations of a traditional RNA ISH method while also providing robust single molecule RNA detection at single cell resolution.

Simultaneous detection of RNA and proteins can reveal cellular sources of secreted proteins, identify specific cell types, and visualize the spatial organization of cells within the tissue. Given the similarities in workflow, the RNAscope ISH assay can be combined with IHC/IF to achieve simultaneous visualization of RNA and protein on the same sample. Referred to as “dual ISH-IHC/IF”, this methodology has been used extensively in neurobiology, immuno-oncology, developmental biology, cell and gene therapy, and inflammation, among other research areas.

In this reference guide are the nearly 300 publications that have used the dual RNAscope ISH-IHC/IF method to simultaneously detect RNA and protein in the same section, categorized by the antigen targeted by IHC/IF in that publication. For more information and recommendations for performing dual ISH-IHC/IF (including technical notes), please visit: www.acdbio.com/dual-ish-and-ihc.

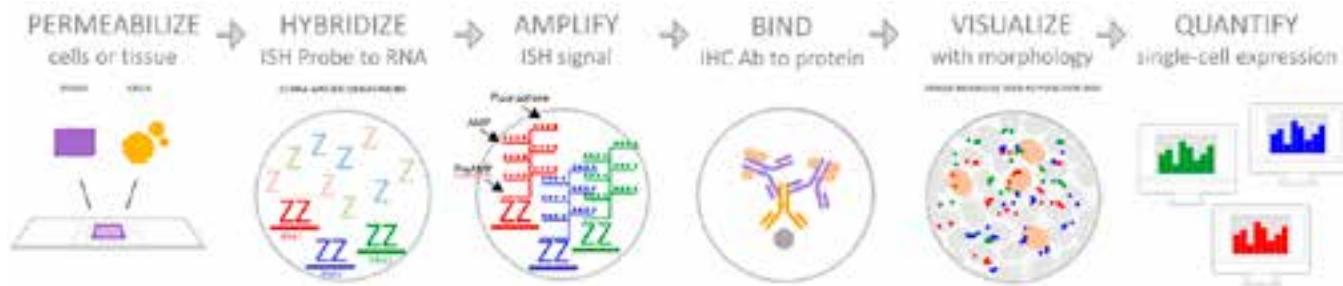


FIGURE 1. General workflow for combining the RNAscope assay with IHC or IF.

Miscellaneous

80HdG	Oligodendroglia are particularly vulnerable to oxidative damage after neurotrauma <i>in vivo</i> Giacci MK, et al. (2018). <i>J Neurosci</i> . DOI: 10.1523/JNEUROSCI.1898-17.2018
AADC (DDC)	Gene therapy for Parkinson's disease: preclinical evaluation of optimally configured TH:CH1 fusion for maximal dopamine synthesis Badin RA, et al. (2019). <i>Mol Ther Methods Clin Dev</i> . DOI: 10.1016/j.omtm.2019.07.002
Annixin 1	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA, et al. (2017). <i>eLife</i> . DOI: 10.7554/eLife.28415
AQP1	WNT5A is transported via lipoprotein particles in the cerebrospinal fluid to regulate hindbrain morphogenesis Kaiser K, et al. (2019). <i>Nat Commun</i> . DOI: 10.1038/s41467-019-09298-4
ARC	A novel environment-evoked transcriptional signature predicts reactivity in single dentate granule neurons Jaeger BN, et al. (2018). <i>Nat Commun</i> . DOI: 10.1038/s41467-018-05418-8
ATPase	Transcellular stomach absorption of a derivatized glucagon-like peptide-1 receptor agonist Buckley ST, et al. (2018). <i>Sci Transl Med</i> . DOI: 10.1126/scitranslmed.aar7047
BAMBI	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA , et al. (2017). <i>eLife</i> . DOI: 10.7554/eLife.28415
BDNF	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA , et al. (2017). <i>eLife</i> . DOI: 10.7554/eLife.28415
Bestrophin-4	Colonic epithelial cell diversity in health and inflammatory bowel disease Parikh K , et al. (2019). <i>Nature</i> . DOI: 10.1038/s41586-019-0992-y
β-gal	Loss of Shh signaling in the neocortex reveals heterogeneous cell recovery responses from distinct oligodendrocyte populations Winkler CC, et al. (2019). <i>Dev Biol</i> . DOI: 10.1016/j.ydbio.2019.04.016
Brn1/2	A translational repression complex in developing mammalian neural stem cells that regulates neuronal specification Zahr SK, et al. (2018). <i>Neuron</i> . DOI: 10.1016/j.neuron.2017.12.045

Miscellaneous (continued)

CA9	Acidification of tumor at stromal boundaries drives transcriptome alterations associated with aggressive phenotypes Rohani N, et al. (2019). <i>Cancer Res.</i> DOI: 10.1158/0008-5472.Can-18-1604
Calretinin	Neuronal heterogeneity and stereotyped connectivity in the auditory afferent system Petitpré C, et al. (2018). <i>Nat Commun.</i> DOI: 10.1038/s41467-018-06033-3
Car2	A Family of non-GPCR chemosensors defines an alternative logic for mammalian olfaction Greer PL, et al. (2016). <i>Cell.</i> DOI: 10.1016/j.cell.2016.05.001
Cav3.2	Detection of single mRNAs in individual cells of the auditory system Salehi P, et al. (2018). <i>Hear Res.</i> DOI: 10.1016/j.heares.2018.07.008
CC1 (APC)	Oligodendroglia are particularly vulnerable to oxidative damage after neurotrauma <i>in vivo</i> Giacci MK, et al. (2018). <i>J Neurosci.</i> DOI: 10.1523/JNEUROSCI.1898-17.2018
CCK	Diversity of interneurons in the dorsal striatum revealed by single-cell RNA sequencing and PatchSeq Muñoz-Manchado AB, et al. (2018). <i>Cell Rep.</i> DOI: 10.1016/j.celrep.2018.07.053
CEL	The mucinous domain of pancreatic carboxyl-ester lipase (CEL) contains core 1/core 2 O-glycans that can be modified by ABO blood group determinants El Jellas K, et al. (2018). <i>J Biol. Chem.</i> DOI: 10.1074/jbc.RA118.001934
CNP	Altered human oligodendrocyte heterogeneity in multiple sclerosis Jäkel S, et al. (2019). <i>Nature.</i> DOI: 10.1038/s41586-019-0903-2
COUPTFII	Immature excitatory neurons develop during adolescence in the human amygdala Sorrells SF, et al. (2019). <i>Nat Commun.</i> DOI: 10.1038/s41467-019-10765-1
CRYAB	Multimodal single-cell analysis reveals physiological maturation in the developing human neocortex Mayer S, et al. (2019). <i>Neuron.</i> DOI: 10.1016/j.neuron.2019.01.027
CTLA4	CTLA-4⁺PD-1⁻ memory CD4⁺ T cells critically contribute to viral persistence in antiretroviral therapy-suppressed, SIV-infected rhesus macaques McGary CS, et al. (2017). <i>Immunity.</i> DOI: 10.1016/j.jimmuni.2017.09.018
Cyp11a1	Zika virus infects human testicular tissue and germ cells Matusali G, et al. (2018). <i>J Clin Invest.</i> DOI: 10.1172/jci121735
DAZL	RNA immunoprecipitation identifies novel targets of DAZL in human foetal ovary Rosario R, et al. (2017). <i>Mol Hum Reprod.</i> DOI: 10.1093/molehr/gax004

Miscellaneous (continued)

DEFA5	Interleukin-2 induces the in vitro maturation of human pluripotent stem cell-derived intestinal organoids Jung KB, et al. (2018). <i>Nat Commun.</i> DOI: 10.1038/s41467-018-05450-8
Desmoplakin	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA , et al. (2017). <i>eLife.</i> DOI: 10.7554/eLife.28415
ERα (ESR1)	Merkel cells activate sensory neural pathways through adrenergic synapses Hoffman BU, et al. (2018). <i>Neuron.</i> DOI: 10.1016/j.neuron.2018.10.034
Fascin (p55)	Lymphatic dissemination of simian immunodeficiency virus after penile inoculation Ma ZM , et al. (2016). <i>J Virol.</i> DOI: 10.1128/jvi.02947-15
FVIII	STAT-3 RNAscope determination in human diffuse large B-cell lymphoma Tamma R, et al. (2019). <i>Transl Oncol.</i> DOI: 10.1016/j.tranon.2018.12.008
G3BP1	Microtubule-driven stress granule dynamics regulate inhibitory immune checkpoint expression in T cells Franchini DM, et al. (2019). <i>Cell Rep.</i> DOI: 10.1016/j.celrep.2018.12.014
G8	Rhabdomyosarcoma and Wilms tumors contain a subpopulation of noggin producing, myogenic cells immunoreactive for lens beaded filament proteins Gerhart J, et al. (2019). <i>PLoS One.</i> DOI: 10.1371/journal.pone.0214758
Gad67	Migrating interneurons secrete fractalkine to promote oligodendrocyte formation in the developing mammalian brain Voronova A, et al. (2017). <i>Neuron.</i> DOI: 10.1016/j.neuron.2017.04.018
GFRα1	SHISA6 confers resistance to differentiation-promoting Wnt/β-Catenin signaling in mouse spermatogenic stem cells Tokue M, et al. (2017). <i>Stem Cell Reports.</i> DOI: 10.1016/j.stemcr.2017.01.006
Glutamine synthetase (Glul)	Inflammasome activation induces pyroptosis in the retina exposed to ocular hypertension injury Pronin A, et al. (2019). <i>Front Mol Neurosci.</i> DOI: 10.3389/fnmol.2019.00036
Glycan 3 (GPC3)	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA , et al. (2017). <i>eLife.</i> DOI: 10.7554/eLife.28415
gp130	Dedifferentiated schwann cell precursors secreting paracrine factors are required for regeneration of the mammalian digit tip Johnston AP, et al. (2016). <i>Cell Stem Cell.</i> DOI: 10.1016/j.stem.2016.06.002

Miscellaneous (continued)

gp38 (Podoplanin)	Colonic CD90+ crypt fibroblasts secrete semaphorins to support epithelial growth Karpus ON, et al. (2019). <i>Cell Rep.</i> DOI: 10.1016/j.celrep.2019.02.101
Gpr37	GPR37 regulates macrophage phagocytosis and resolution of inflammatory pain Bang S, et al. (2018). <i>J Clin Invest.</i> DOI: 10.1172/jci99888
GS Lectin AlexaFluor 488	Simultaneous visualization and cell-specific confirmation of RNA and protein in the mouse retina Stempel AJ, et al. (2014). <i>Mol Vis.</i> PMID: 25352743
GZMB	A case report of clonal EBV-like memory CD4+ T cell activation in fatal checkpoint inhibitor-induced encephalitis Johnson DB, et al. (2019). <i>Nat Med.</i> DOI: 10.1038/s41591-019-0523-2
HCRT	Involvement of PLAGL1/ZAC1 in hypocretin/orexin transcription Tanaka S, et al. (2019). <i>Int J Mol Med.</i> DOI: 10.3892/ijmm.2019.4143
HERV (Syncytin, ERVW1)	Dengue virus immunity increases Zika virus-induced damage during pregnancy Brown JA, et al. (2019). <i>Immunity.</i> DOI: 10.1016/j.jimmuni.2019.01.005
HLA-DR	Tumor-specific MHC-II expression drives a unique pattern of resistance to immunotherapy via LAG-3/FCRL6 engagement Johnson DB, et al. (2018). <i>JCI Insight.</i> DOI: 10.1172/jci.insight.120360
HNF4a	H19 is expressed in hybrid hepatocyte nuclear factor 4α⁺ periportal hepatocytes but not cytokeratin 19⁺ cholangiocytes in cholestatic livers Jiang Y, et al. (2018). <i>Hepatol Commun.</i> DOI: 10.1002/hep4.1252
HOPX	Multimodal single-cell analysis reveals physiological maturation in the developing human neocortex Mayer S, et al. (2019). <i>Neuron.</i> DOI: 10.1016/j.neuron.2019.01.027
HSD2	Aldosterone-sensitive HSD2 neurons in mice Gasparini S, et al. (2019). <i>Brain Struct Funct.</i> DOI: 10.1007/s00429-018-1778-y
HTII-280	MicroRNA-29c prevents pulmonary fibrosis by regulating epithelial cell renewal and apoptosis Xie T, et al. (2017). <i>Am J Respir Cell Mol Biol.</i> DOI: 10.1165/rcmb.2017-0133OC
Int	Oxytocin/vasopressin-like peptide inotocin regulates cuticular hydrocarbon synthesis and water balancing in ants Koto A, et al. (2019). <i>Proc Natl Acad Sci U S A.</i> DOI: 10.1073/pnas.1817788116
IRF8	GPR34 in spinal microglia exacerbates neuropathic pain in mice Sayo A, et al. (2019). <i>J Neuroinflammation.</i> DOI: 10.1186/s12974-019-1458-8

Miscellaneous (continued)

Isl1/2	Motor neurons control blood vessel patterning in the developing spinal cord Himmels P, et al. (2017). <i>Nat Commun.</i> DOI: 10.1038/ncomms14583
KAZALD1	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA , et al. (2017). <i>eLife.</i> DOI: 10.7554/eLife.28415
Laminin	Netrin-1 confines rhombic lip-derived neurons to the CNS Yung AR, et al. (2018). <i>Cell Rep.</i> DOI: 10.1016/j.celrep.2018.01.068
Langerin	Identification of HIV transmitting CD11c⁺ human epidermal dendritic cells Bertram KM, et al. (2019). <i>Nat Commun.</i> DOI: 10.1038/s41467-019-10697-w
LAMP1	Senolytic therapy alleviates Aβ-associated oligodendrocyte progenitor cell senescence and cognitive deficits in an Alzheimer's disease model Zhang P, et al. (2019). <i>Nat Neurosci.</i> DOI: 10.1038/s41593-019-0372-9
LOR	Single-cell transcriptomics reveals that differentiation and spatial signatures shape epidermal and hair follicle heterogeneity Joost S, et al. (2016). <i>Cell Syst.</i> DOI: 10.1016/j.cels.2016.08.010
MAC-2 (LGALS3)	Dengue virus immunity increases Zika virus-induced damage during pregnancy Brown JA, et al. (2019). <i>Immunity.</i> DOI: 10.1016/j.immuni.2019.01.005

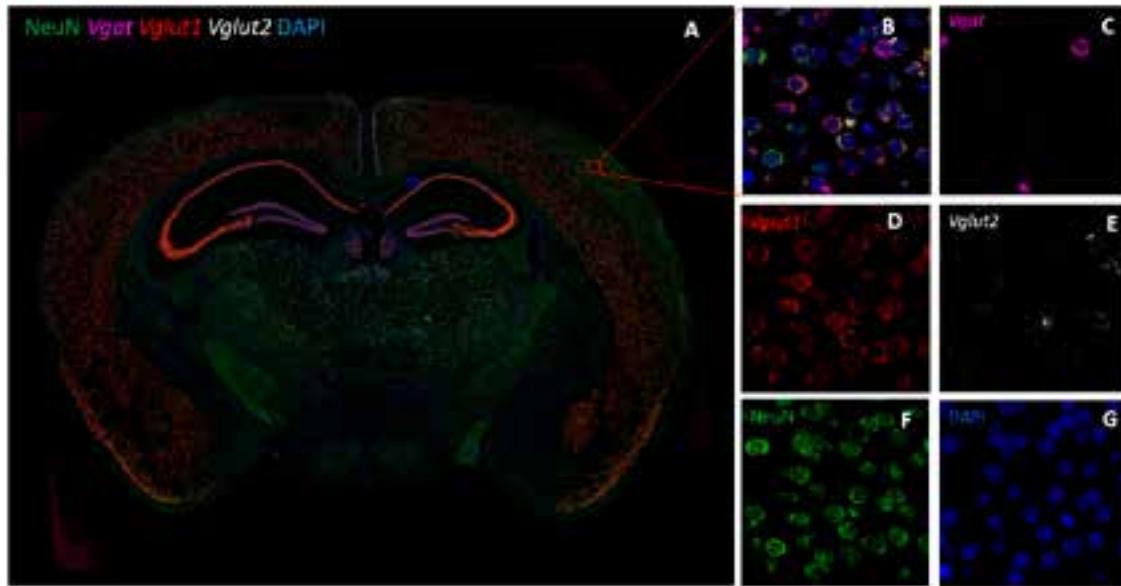


FIGURE 2. Spatial mapping of glutamatergic and GABAergic neurons in the mouse brain. The RNAscope Multiplex Fluorescent V2 Assay for Vgat, Vglut1, and Vglut2 was combined with IF for NeuN.

Miscellaneous (continued)

MAP2	ALS along the axons – expression of coding and noncoding RNA differs in axons of ALS models Rotem N, et al. (2017). <i>Sci Rep.</i> DOI: 10.1038/srep44500
MCT	Possible involvement of human mast cells in the establishment of pregnancy via killer cell Ig-like receptor 2DL4 Ueshima C, et al. (2018). <i>Am J Pathol.</i> DOI: 10.1016/j.ajpath.2018.02.012
MEF2C	Dusp6 attenuates Ras/MAPK signaling to limit zebrafish heart regeneration Missinato MA, et al. (2018). <i>Development.</i> DOI: 10.1242/dev.157206
MERTK	Apoptotic tumor cell-derived microRNA-375 uses CD36 to alter the tumor-associated macrophage phenotype Frank AC, et al. (2019). <i>Nat Commun.</i> DOI: 10.1038/s41467-019-08989-2
MPO	Single-cell transcriptomics of human and mouse lung cancers reveals conserved myeloid populations across individuals and species Zilionis R, et al. (2019). <i>Immunity.</i> DOI: 10.1016/j.jimmuni.2019.03.009
MYRF	Altered human oligodendrocyte heterogeneity in multiple sclerosis Jäkel S, et al. (2019). <i>Nature.</i> DOI: 10.1038/s41586-019-0903-2
Na,K-ATPase (ATA1A3)	Detection of single mRNAs in individual cells of the auditory system Salehi P, et al. (2018) <i>Hear Res.</i> DOI: 10.1016/j.heares.2018.07.008
Nephrin	Dysregulated expression but redundant function of the long non-coding RNA HOTAIR in diabetic kidney disease Majumder S, et al. (2019). <i>Diabetologia.</i> DOI: 10.1007/s00125-019-4967-1
Nexilin	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA, et al. (2017). <i>eLife.</i> DOI: 10.7554/eLife.28415
Nidogen-2	Spatially and functionally distinct subclasses of breast cancer-associated fibroblasts revealed by single cell RNA sequencing Bartoschek M, et al. (2018). <i>Nat Commun.</i> DOI: 10.1038/s41467-018-07582-3
Nkx2.2	Glial- and neuronal-specific expression of CCL5 mRNA in the rat brain Lanfranco MF, et al. (2018). <i>Front Neuroanat.</i> DOI: 10.3389/fnana.2017.00137
Notch1	A method for manual and automated multiplex RNAscope <i>in situ</i> hybridization and immunocytochemistry on cytospin samples Chan S, et al. (2018). <i>PLoS One.</i> DOI: 10.1371/journal.pone.0207619

Miscellaneous (continued)

OLFM4	Interleukin-2 induces the in vitro maturation of human pluripotent stem cell-derived intestinal organoids Jung KB, et al. (2018). <i>Nat Commun.</i> DOI: 10.1038/s41467-018-05450-8
OXTR	Peri-infarct upregulation of the oxytocin receptor in vascular dementia McKay EC, et al. (2019). <i>J Neuropathol Exp Neurol.</i> DOI: 10.1093/jnen/nlz023
p24	Elucidating the burden of HIV in tissues using multiplexed immunofluorescence and <i>in situ</i> hybridization: methods for the single-cell phenotypic characterization of cells harboring HIV <i>in situ</i> Vasquez JJ, et al. (2018). <i>J Histochem Cytochem.</i> DOI: 10.1369/0022155418756848
P2Y12 (P2RY12, Purinergic Receptor)	Sexual dimorphism in the inflammatory response to traumatic brain injury Villapol S, et al. (2017). <i>Glia.</i> DOI: 10.1002/glia.23171
P75ntr (NGFR)	Morphological characteristics of p75 neurotrophin receptor-positive cells define a new type of glial cell in the rat dorsal root ganglia Koike T, et al. (2019). <i>J Comp Neurol.</i> DOI: 10.1002/cne.24667
Parvalbumin	Sensory neuron diversity in the inner ear is shaped by activity Shrestha BR, et al. (2018). <i>Cell.</i> DOI: 10.1016/j.cell.2018.07.007
PDF	Transcriptional regulation of lipophorin receptors supports neuronal adaptation to chronic elevations of activity Yin J, et al. (2018). <i>Cell Rep.</i> DOI: 10.1016/j.celrep.2018.10.016
PERK	Central relaxin-3 receptor (RXFP3) activation impairs social recognition and modulates ERK phosphorylation in specific GABAergic amygdala neurons Albert-Gasco H, et al. (2019). <i>Brain Struct Funct.</i> DOI: 10.1007/s00429-018-1763-5
PgR (Progesterone Receptor)	Oestrogen receptor α AF-1 and AF-2 domains have cell population-specific functions in the mammary epithelium Cagnet S, et al. (2018). <i>Nat Commun.</i> DOI: 10.1038/s41467-018-07175-0
PKP2	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA , et al. (2017). <i>eLife.</i> DOI: 10.7554/eLife.28415
PLAGL1	Involvement of PLAGL1/ZAC1 in hypocretin/orexin transcription Tanaka S, et al. (2019). <i>Int J Mol Med.</i> DOI: 10.3892/ijmm.2019.4143
PLEKHA7	Cadherin complexes recruit mRNAs and RISC to regulate epithelial cell signaling Kourtidis A, et al. (2017). <i>J Cell Biol.</i> DOI: 10.1083/jcb.201612125

Miscellaneous (continued)

Plin2	Obesity-induced cellular senescence drives anxiety and impairs neurogenesis Ogrodnik M, et al. (2019). <i>Cell Metab.</i> DOI: 10.1016/j.cmet.2018.12.008
PPP1R17	Multimodal single-cell analysis reveals physiological maturation in the developing human neocortex Mayer S, et al. (2019). <i>Neuron.</i> DOI: 10.1016/j.neuron.2019.01.027
Ppy	Use of the Fluidigm C1 platform for RNA sequencing of single mouse pancreatic islet cells Xin Y, et al. (2016). <i>Proc Natl Acad Sci U S A.</i> DOI: 10.1073/pnas.1602306113
pro-MBP1 (PRG2)	Basophil recruitment into tumor-draining lymph nodes correlates with Th2 inflammation and reduced survival in pancreatic cancer patients De Monte L, et al. (2016). <i>Cancer Res.</i> DOI: 10.1158/0008-5472.can-15-1801-t
Prospc (Pro-surfactant Protein C)	The strength of mechanical forces determines the differentiation of alveolar epithelial cells Li J, et al. (2018). <i>Dev Cell.</i> DOI: 10.1016/j.devcel.2018.01.008
PrRP (PRLH)	GLP-1 neurons form a local synaptic circuit within the rodent nucleus of the solitary tract Card JP, et al. (2018). <i>J Comp Neurol.</i> DOI: 10.1002/cne.24482
pSHP-1	PD-L1 inhibits acute and chronic pain by suppressing nociceptive neuron activity via PD-1 Chen G, et al. (2017). <i>Nat Neurosci.</i> DOI 10.1038/nn.4571
pSTAT3	Brs3 neurons in the mouse dorsomedial hypothalamus regulate body temperature, energy expenditure, and heart rate, but not food intake Piñol RA, et al. (2018). <i>Nat Neurosci.</i> DOI: 10.1038/s41593-018-0249-3
PTBP3	LncRNA Meg3 protects endothelial function by regulating the DNA damage response Shihabudeen Haider Ali MS, et al. (2019). <i>Nucleic Acids Res.</i> DOI: 10.1093/nar/gky1190
PTEN	Cabozantinib eradicates advanced murine prostate cancer by activating antitumor innate immunity Patnaik A, et al. (2017). <i>Cancer Discov.</i> DOI: 10.1158/2159-8290.cd-16-0778
RBPMS	Inflammasome activation induces pyroptosis in the retina exposed to ocular hypertension injury Pronin A, et al. (2019). <i>Front Mol Neurosci.</i> DOI: 10.3389/fnmol.2019.00036
Renin	Angiotensin II short-loop feedback: Is there a role of Ang II for the regulation of the renin system <i>in vivo</i>? Neubauer B, et al. (2018). <i>Hypertension.</i> DOI: 10.1161/hypertensionaha.117.10357

Miscellaneous (continued)

Rh6	The HisCl1 histamine receptor acts in photoreceptors to synchronize Drosophila behavioral rhythms with light-dark cycles Alejevski F, et al. (2019). <i>Nat Commun.</i> DOI: 10.1038/s41467-018-08116-7
SAHH	Combined single-cell profiling of lncRNAs and functional screening reveals that H19 is pivotal for embryonic hematopoietic stem cell development Zhou J, et al. (2019). <i>Cell Stem Cell.</i> DOI: 10.1016/j.stem.2018.11.023
SFTPC	Expansion of hedgehog disrupts mesenchymal identity and induces emphysema phenotype Wang C, et al. (2018). <i>J Clin Invest.</i> DOI: 10.1172/jci99435
SM22alpha (Tagln)	Feasibility evaluation of myocardial cannabinoid type 1 receptor imaging in obesity: a translational approach Valenta I, et al. (2018). <i>JACC Cardiovasc Imaging.</i> DOI: 10.1016/j.jcmg.2017.11.019
Smaug2 (SAMD4B)	A Smaug2-based translational repression complex determines the balance between precursor maintenance versus differentiation during mammalian neurogenesis Amadei G, et al. (2015). <i>J Neurosci.</i> DOI: 10.1523/jneurosci.2172-15.2015
SSEA-1 (CD15, FUT4)	Does human endometrial LGR5 gene expression suggest the existence of another hormonally regulated epithelial stem cell niche? Tempest N, et al. (2018). <i>Hum Reprod.</i> DOI: 10.1093/humrep/dey083
STAR	Zika virus infects human testicular tissue and germ cells Matusali G, et al. (2018). <i>J Clin Invest.</i> DOI: 10.1172/jci121735
Synaptotagmin	The local expression and trafficking of tyrosine hydroxylase mRNA in the axons of sympathetic neurons Gervasi NM, et al. (2016). <i>RNA.</i> DOI: 10.1261/rna.053272.115
Tesclacin	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA, et al. (2017). <i>eLife.</i> DOI: 10.7554/eLife.28415
TFPI2	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA, et al. (2017). <i>eLife.</i> DOI: 10.7554/eLife.28415
TGFβ	Inhibition of Nox4-dependent ROS signaling attenuates prostate fibroblast activation and abrogates stromal-mediated protumorigenic interactions Sampson N, et al. (2018). <i>Int J Cancer.</i> DOI: 10.1002/ijc.31316

Miscellaneous (continued)

TOMM20	CLUH regulates mitochondrial metabolism by controlling translation and decay of target mRNAs Schatton D, et al. (2017). <i>J Cell Biol.</i> DOI: 10.1083/jcb.201607019
TRAcP (ACP5)	Coupling of bone resorption and formation in real time: new knowledge gained from human haversian BMUs Lassen NE, et al. (2017). <i>JJ Bone Miner Res.</i> DOI: 10.1002/jbmr.3091
Tropomyosin	Myocardial polyploidization creates a barrier to heart regeneration in zebrafish González-Rosa JM, et al. (2018). <i>Dev Cell.</i> DOI: 10.1016/j.devcel.2018.01.021
Troponin I1	Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions Morrison JA, et al. (2017). <i>eLife.</i> DOI: 10.7554/eLife.28415
TSLP	A single-cell survey of the small intestinal epithelium Haber AL, et al. (2017). <i>Nature.</i> DOI: 10.1038/nature24489
Ubiquitin	High-dimensional cell-level analysis of tissues with Ce3D multiplex volume imaging Li W, et al. (2019). <i>Nat Protoc.</i> DOI: 10.1038/s41596-019-0156-4
VCAM1	Developmental emergence of adult neural stem cells as revealed by single-cell transcriptional profiling Yuzwa SA, et al. (2017). <i>Cell Rep.</i> DOI: 10.1016/j.celrep.2017.12.017
VDR	Vitamin D differentially regulates colon stem cells in patient-derived normal and tumor organoids Fernández-Barral A, et al. (2019). <i>FEBS J.</i> DOI: 10.1111/febs.14998
VEGFR3	Transcriptional landscape of pulmonary lymphatic endothelial cells during fetal gestation Norman TA Jr, et al. (2019). <i>PLoS One.</i> DOI: 10.1371/journal.pone.0216795
YBX1	Satellite RNAs promote pancreatic oncogenic processes via the dysfunction of YBX1 Kishikawa T, et al. (2016). <i>Nat Commun.</i> DOI: 10.1038/ncomms13006
YTHDF2	Suppression of m⁶A reader Ythdf2 promotes hematopoietic stem cell expansion Li Z, et al. (2018). <i>Cell Res.</i> DOI: 10.1038/s41422-018-0072-0

Virus-Related Antibodies

Anti-ZIKV NS3	Zika virus production is resistant to RNase L antiviral activity Whelan JN, et al. (2019). <i>J Virol.</i> DOI: 10.1128/jvi.00313-19
dsRNA	A novel sheet-like virus particle array is a hallmark of Zika virus infection Liu J, et al. (2018). <i>Emerg Microbes Infect.</i> DOI: 10.1038/s41426-018-0071-8
Flavivirus	A novel sheet-like virus particle array is a hallmark of Zika virus infection Liu J, et al. (2018). <i>Emerg Microbes Infect.</i> DOI: 10.1038/s41426-018-0071-8
gD (LP14)	Multiple posttranscriptional strategies to regulate the herpes simplex virus 1 vhs endoribonuclease Elliott G, et al. (2018). <i>J Virol.</i> DOI: 10.1128/jvi.00818-18
HIV p24/GAG	Visualization of HIV-1 RNA transcription from integrated HIV-1 DNA in reactivated latently infected cells Ukah OB, et al. (2018). <i>Viruses.</i> DOI: 10.3390/v10100534
HIV-1 p24	Inhibition of heat shock protein 90 prevents HIV rebound Joshi P, et al. (2016). <i>J Biol Chem.</i> DOI: 10.1074/jbc.M116.717538
HIVp24	Cellular HIV reservoirs and viral rebound from the lymphoid compartments of 4'-ethynyl-2-fluoro-2'-deoxyadenosine (EFdA)-suppressed humanized mice Maidji E, et al. (2019). <i>Viruses.</i> DOI: 10.3390/v11030256
LCMV	Neurons under T cell attack coordinate phagocyte-mediated synaptic stripping Di Liberto G, et al. (2018). <i>Cell.</i> DOI: 10.1016/j.cell.2018.07.049
PyMT	Loss of amphiregulin reduces myoepithelial cell coverage of mammary ducts and alters breast tumor growth Mao SPH, et al. (2018). <i>Breast Cancer Res.</i> DOI: 10.1186/s13058-018-1057-0
V5-vhs	Multiple posttranscriptional strategies to regulate the herpes simplex virus 1 vhs endoribonuclease Elliott G, et al. (2018). <i>J Virol.</i> DOI: 10.1128/jvi.00818-18

Beta amyloid

Senolytic therapy alleviates Aβ-associated oligodendrocyte progenitor cell senescence and cognitive deficits in an Alzheimer's disease model Zhang P, et al. (2019). <i>Nat Neurosci.</i> DOI: 10.1038/s41593-019-0372-9
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Human Alzheimer's disease gene expression signatures and immune profile in APP mouse models: a discrete transcriptomic view of A β plaque pathology

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PHD2 inactivation in Type I cells drives HIF-2 α -dependent multilineage hyperplasia and the formation of paraganglioma-like carotid bodies

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Colonic CD90+ crypt fibroblasts secrete semaphorins to support epithelial growth

Karpus ON, et al. (2019). *Cell Rep*. DOI: 10.1016/j.celrep.2019.02.101

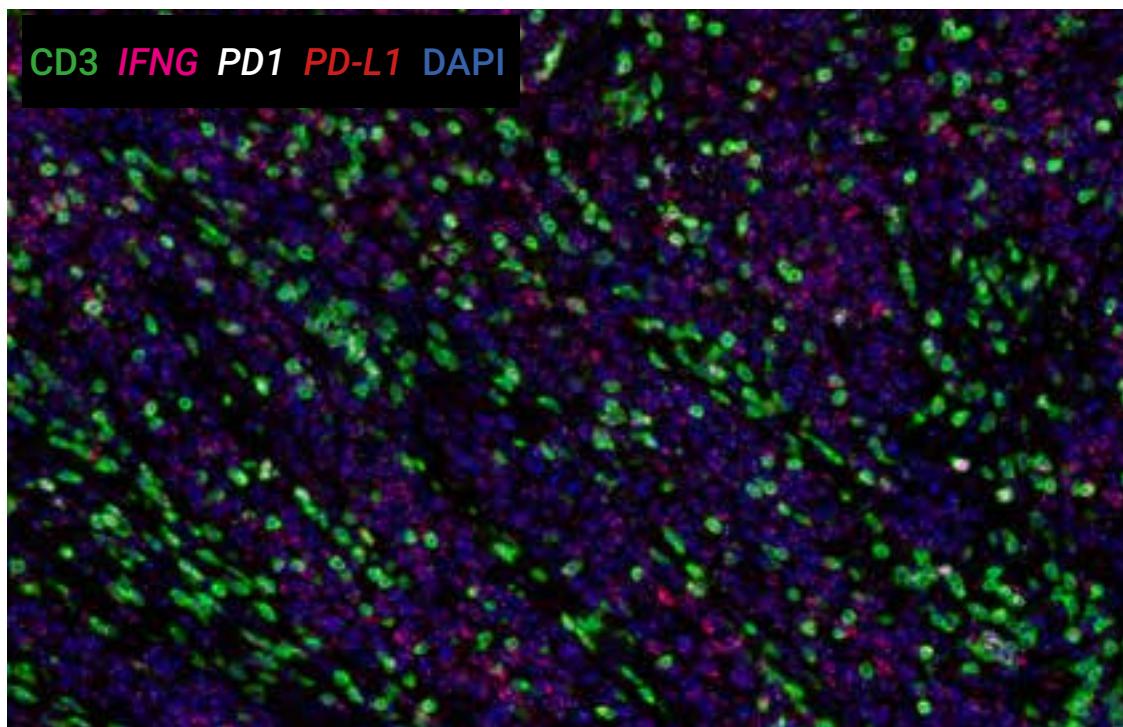


FIGURE 3. Immune cell characterization in the tumor microenvironment. The RNAscope Multiplex Fluorescent V2 Assay for IFNG, PD1, and PD-L1 was combined with IF for CD3.

Cadherins

Reference component analysis of single-cell transcriptomes elucidates cellular heterogeneity in human colorectal tumors

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Cadherin complexes recruit mRNAs and RISC to regulate epithelial cell signaling

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CD8⁺ T cells modulate autosomal dominant polycystic kidney disease progression

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T helper cell cytokines modulate intestinal stem cell renewal and differentiation

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Self-maintaining gut macrophages are essential for intestinal homeostasis

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Tumour-vasculature development via endothelial-to-mesenchymal transition after radiotherapy controls CD44v6⁺ cancer cell and macrophage polarization

Choi SH, et al. (2018). *Nat Commun*. DOI: 10.1038/s41467-018-07470-w

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CSF-1 controls cerebellar microglia and is required for motor function and social interaction

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Mechanically activated Piezo channels mediate touch and suppress acute mechanical pain response in mice

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Lymphatic dissemination of simian immunodeficiency virus after penile inoculation

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Wildenberg ME, et al. (2017). *J Crohns Colitis.* DOI: 10.1093/ecco-jcc/jjx104

Liver macrophage-associated inflammation correlates with SIV burden and is substantially reduced following cART

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Accumulation of follicular CD8+ T cells in pathogenic SIV infection

Ferrando-Martinez S, et al. (2018). *J Clin Invest.* DOI: 10.1172/jci96207

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CD32-RNA Co-localizes with HIV-RNA in CD3+ Cells Found within Gut Tissues from Viremic and ART-Suppressed Individuals

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CD4

The presence of interleukin-13 at pancreatic ADM/PanIN lesions alters macrophage populations and mediates pancreatic tumorigenesis

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Pomalidomide alters pancreatic macrophage populations to generate an immune-responsive environment at precancerous and cancerous lesions

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CD8

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CD11

Epigenetic promoter DNA methylation of miR-124 promotes HIV-1 Tat-mediated microglial activation via MECP2-STAT3 axis

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Accumulation of worn-out GBM material substantially contributes to mesangial matrix expansion in diabetic nephropathy

Kriz W, et al. (2017). *Am J Physiol Renal Physiol*. DOI: 10.1152/ajprenal.00020.2017

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The proton-activated receptor GPR4 modulates intestinal inflammation

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Single-cell transcriptomics reveals that differentiation and spatial signatures shape epidermal and hair follicle heterogeneity

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Integrated Human Evaluation of the Lysophosphatidic Acid Pathway as a Novel Therapeutic Target in Atherosclerosis

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VEGF receptor-2/neuropilin 1 trans-complex formation between endothelial and tumor cells is an independent predictor of pancreatic cancer survival

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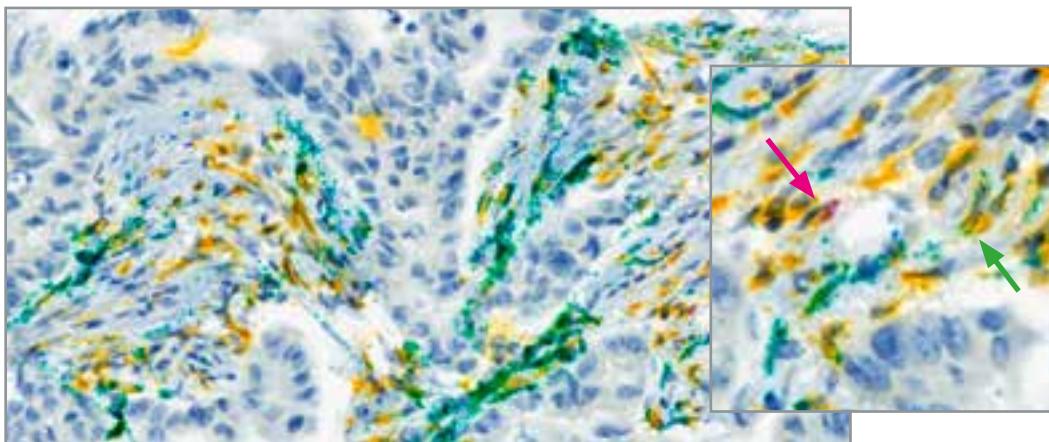


FIGURE 4. Detection of activated macrophages in the TME of human lung cancer. The RNAscope VS Duplex Assay for *IL-12* and *CXCL9* was combined with IHC for CD68.

CD45

Phosphatidylinositol 3-kinase delta pathway: a novel therapeutic target for Sjögren's syndrome

Nayar S, et al. (2019). *Ann Rheum Dis*. DOI: 10.1136/annrheumdis-2017-212619

Recurrent PDGFRB mutations in unicentric Castleman disease

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CD68

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Reactivation of simian immunodeficiency virus reservoirs in the brain of virally suppressed macaques

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IL-12 ISH
CXCL9 ISH
CD68 IHC

CD68 (continued)

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The meningeal lymphatic system: a route for HIV brain migration?

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Live imaging of HIV-1 transfer across T cell virological synapse to epithelial cells that promotes stromal macrophage infection

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Zika virus infects human testicular tissue and germ cells

Matusali G, et al. (2018). *J Clin Invest.* DOI: 10.1172/jci121735

Macrophages but not astrocytes harbor HIV DNA in the brains of HIV-1-infected aviremic individuals on suppressive antiretroviral therapy

Ko A, et al. (2019). *J Neuroimmune Pharmacol.* DOI: 10.1007/s11481-018-9809-2

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CD163

Replication of CMV in the gut of HIV-infected individuals and epithelial barrier dysfunction

Maidji E, et al. (2017). *PLoS Pathog.* DOI: 10.1371/journal.ppat.1006202

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Zika virus infects human testicular tissue and germ cells

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Human tumor-associated macrophage and monocyte transcriptional landscapes reveal cancer-specific reprogramming, biomarkers, and therapeutic targets

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Other CD Markers

Pericyte loss leads to circulatory failure and pleiotrophin depletion causing neuron loss

Nikolakopoulou AM, et al. (2019). *Nat Neurosci.* DOI: 10.1038/s41593-019-0434-z

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A method for manual and automated multiplex RNAscope *in situ* hybridization and immunocytochemistry on cytospin samples

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Other CD Markers (*continued*)

Reduced lymphotoxin-beta production by tumour cells is associated with loss of follicular dendritic cell phenotype and diffuse growth in follicular lymphoma

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Dawes JM, et al. (2018). *Neuron.* DOI: 10.1016/j.neuron.2018.01.033

Comprehensive analysis of long noncoding RNA expression in dorsal root ganglion reveals cell-type specificity and dysregulation after nerve injury

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Mechanically activated Piezo channels mediate touch and suppress acute mechanical pain response in mice

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Lineage dynamics of murine pancreatic development at single-cell resolution

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Cytokeratins

Co-detection of miR-21 and TNF- α mRNA in budding cancer cells in colorectal cancer

Møller T, et al. (2019). *Int J Mol Sci*. DOI 10.3390/ijms20081907

Establishment of EBV latency in nasopharyngeal tumor epithelial cells by in vivo cell-mediated transfer infection

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H19 Is expressed in hybrid hepatocyte nuclear factor 4 α^+ periportal hepatocytes but not cytokeratin 19 $^+$ cholangiocytes in cholestatic livers

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Interleukin-6 blockade attenuates lung cancer tissue construction integrated by cancer stem cells

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Brown JA, et al. (2019). *Immunity.* DOI: 10.1016/j.jimmuni.2019.01.005

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Targeting LIF-mediated paracrine interaction for pancreatic cancer therapy and monitoring

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Characterization of a BAC transgenic mouse expressing Krt19-driven iCre recombinase in its digestive organs

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Single-cell analysis identifies LY6D as a marker linking castration-resistant prostate luminal cells to prostate progenitors and cancer

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Increased lactate secretion by cancer cells sustains non-cell-autonomous adaptive resistance to MET and EGFR targeted therapies

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Single-cell analysis of the liver epithelium reveals dynamic heterogeneity and an essential role for YAP in homeostasis and regeneration

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DCLK1

The presence of interleukin-13 at pancreatic ADM/PanIN lesions alters macrophage populations and mediates pancreatic tumorigenesis

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Microtubule-driven stress granule dynamics regulate inhibitory immune checkpoint expression in T cells

Franchini DM, et al. (2019). *Cell Rep*. DOI: 10.1016/j.celrep.2018.12.014

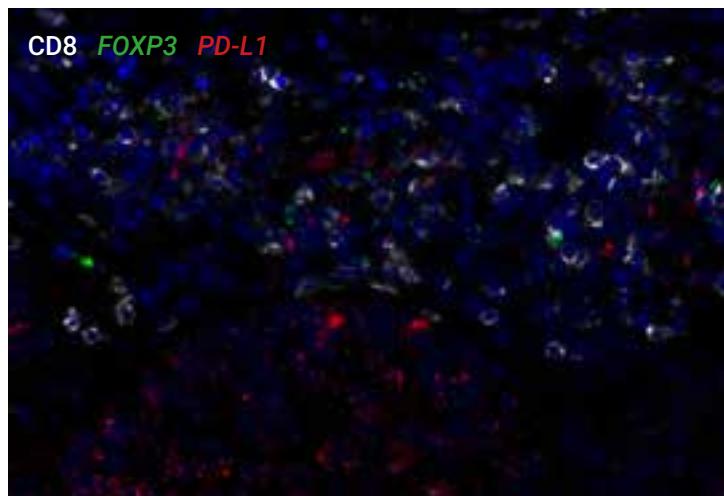


FIGURE 5. Characterization of immune cell infiltration into the TME in human lung cancer. The RNAscope Multiplex Fluorescent V2 Assay for *FOXP3* and *PD-L1* was combined with IF for CD3.

DCX

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Immature excitatory neurons develop during adolescence in the human amygdala

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DDX4

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H19 is expressed in hybrid hepatocyte nuclear factor 4 α^+ periportal hepatocytes but not cytokeratin 19 $^+$ cholangiocytes in cholestatic livers

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dsRed

Melanin-concentrating hormone neurons promote rapid eye movement sleep independent of glutamate release

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F4/80 (continued)

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Cellular localization and regulation of receptors and enzymes of the endocannabinoid system in intestinal and systemic inflammation

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GFAP

Simultaneous visualization and cell-specific confirmation of RNA and protein in the mouse retina

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Translation in astrocyte distal processes sets molecular heterogeneity at the gliovascular interface

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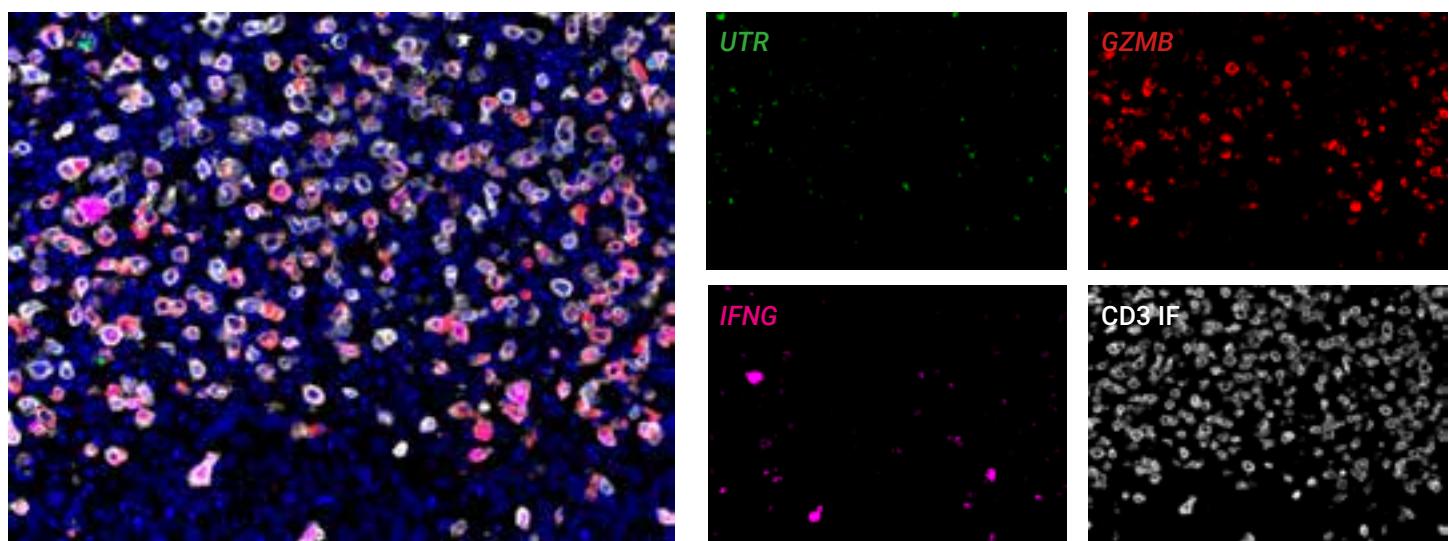


FIGURE 6. Visualization of activated CAR-T cells in human xenograft tumor. The RNAscope Multiplex Fluorescent V2 Assay for *IFNG*, *GZMB*, and the 3' UTR of a CAR vector was combined with IF for *CD3*.

GFAP (continued)

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New role of P2X7 receptor in an Alzheimer's disease mouse model

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Single cell RNA sequencing identifies TGF β as a key regenerative cue following LPS-induced lung injury

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Glucagon

GPR142 prompts glucagon-like Peptide-1 release from islets to improve β cell function

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GPR119 agonism increases glucagon secretion during insulin-induced hypoglycemia

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Glut1

FZD₁₀-Gα_{i3} signalling axis points to a role of FZD₁₀ in CNS angiogenesis

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Fibronectin produced by cerebral endothelial and vascular smooth muscle cells contributes to perivascular extracellular matrix in late-delayed radiation-induced brain injury

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HNK1

Resolving *in vivo* gene expression during collective cell migration using an integrated RNAscope, immunohistochemistry and tissue clearing method

Morrison JA, et al. (2017). *Mech Dev.* DOI: 10.1016/j.mod.2017.06.004

Single-cell transcriptome analysis of avian neural crest migration reveals signatures of invasion and molecular transitions

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IB4

Motor neurons control blood vessel patterning in the developing spinal cord

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Immune or genetic-mediated disruption of CASPR2 causes pain hypersensitivity due to enhanced primary afferent excitability

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Translation in astrocyte distal processes sets molecular heterogeneity at the gliovacular interface

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A critical role for dopamine D5 receptors in pain chronicity in male mice

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Sustained inflammation after pericyte depletion induces irreversible blood-retina barrier breakdown

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NOX2 deficiency alters macrophage phenotype through an IL-10/STAT3 dependent mechanism: implications for traumatic brain injury

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Migrating interneurons secrete fractalkine to promote oligodendrocyte formation in the developing mammalian brain

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Sexual dimorphism in the inflammatory response to traumatic brain injury

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Rationally engineered AAV capsids improve transduction and volumetric spread in the CNS

Kanaan NM, et al. (2017). *Mol Ther Nucleic Acids*. DOI: 10.1016/j.omtn.2017.06.011

HIV-infected macrophages and microglia that survive acute infection become viral reservoirs by a mechanism involving Bim

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Increased microglial CSF1R expression in the SIV/Macaque model of HIV CNS disease

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Glial- and neuronal-specific expression of CCL5 mRNA in the rat brain

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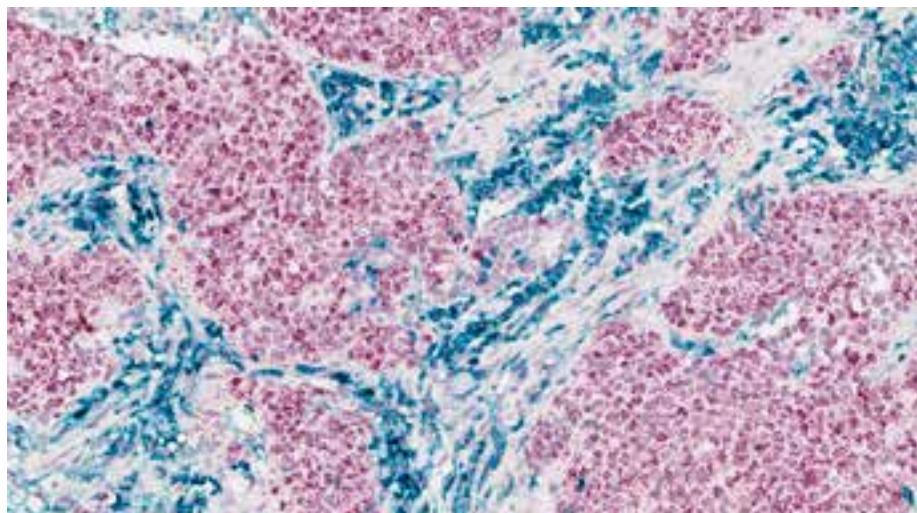


FIGURE 7. **Discerning tumor and stromal gene expression.** The RNAscope 2.5 HD Assay-Red for *PD-L1* was combined with IHC for CD45.

Iba1 (continued)

New role of P2X7 receptor in an Alzheimer's disease mouse model

Martin E, et al. (2019). *Mol Psychiatry*. DOI: 10.1038/s41380-018-0108-3

Simian immunodeficiency virus persistence in cellular and anatomic reservoirs in antiretroviral therapy-suppressed infant rhesus macaques

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Macrophages regulate schwann cell maturation after nerve injury

Stratton JA, et al. (2018). *Cell Rep*. DOI: 10.1016/j.celrep.2018.08.004

PDGFR β cells rapidly relay inflammatory signal from the circulatory system to neurons via chemokine CCL2

Duan L, et al. (2018). *Neuron*. DOI: 10.1016/j.neuron.2018.08.030

Alzheimer's disease phospholipase C-gamma-2 (PLCG2) protective variant is a functional hypermorph

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Inflammasome activation induces pyroptosis in the retina exposed to ocular hypertension injury

Pronin A, et al. (2019). *Front Mol Neurosci*. DOI: 10.3389/fnmol.2019.00036

C3- and CR3-dependent microglial clearance protects photoreceptors in retinitis pigmentosa

Silverman SM, et al. (2019). *J Exp Med*. DOI: 10.1084/jem.20190009

Silencing alpha synuclein in mature nigral neurons results in rapid neuroinflammation and subsequent toxicity

Benskey MJ, et al. (2018). *Front Mol Neurosci*. DOI: 10.3389/fnmol.2018.00036

A combination of ontogeny and CNS environment establishes microglial identity

Bennett FC, et al. (2018). *Neuron*. DOI: 10.1016/j.neuron.2018.05.014

Combination of fluorescent *in situ* hybridization (FISH) and immunofluorescence imaging for detection of cytokine expression in microglia/macrophage Cells

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Ig Molecules

Single-cell RNA sequencing of lymph node stromal cells reveals niche-associated heterogeneity

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S-Geranylgeranyl-L-glutathione is a ligand for human B cell-confinement receptor P2RY8

Lu E, et al. (2019). *Nature*. DOI: 10.1038/s41586-019-1003-z

A method for manual and automated multiplex RNAscope *in situ* hybridization and immunocytochemistry on cytocentrifuge samples

Chan S, et al. (2018). *PLoS One*. DOI: 10.1371/journal.pone.0207619

Insulin

Suppression of insulin production and secretion by a decretin hormone

Alfa RW, et al. (2015). *Cell Metab*. DOI: 10.1016/j.cmet.2015.01.006

GPR142 prompts glucagon-like Peptide-1 release from islets to improve β cell function

Lin HV, et al. (2018). *Mol Metab*. DOI: 10.1016/j.molmet.2018.02.008

GPR119 agonism increases glucagon secretion during insulin-induced hypoglycemia

Li NX, et al. (2018). *Diabetes*. DOI: 10.2337/db18-0031

Use of the Fluidigm C1 platform for RNA sequencing of single mouse pancreatic islet cells

Xin Y, et al. (2016). *Proc Natl Acad Sci U S A*. DOI: 10.1073/pnas.1602306113

Ki67

Distribution of Lgr5-positive cancer cells in intramucosal gastric signet-ring cell carcinoma

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Multimodal single-cell analysis reveals physiological maturation in the developing human neocortex

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A case report of clonal EBV-like memory CD4+ T cell activation in fatal checkpoint inhibitor-induced encephalitis

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Lysozyme

Automated double *in situ* detection of mouse Lgr5 mRNA and lysozyme protein in examining the neighboring cell types of the mouse intestinal crypt

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Radical and lunatic fringes modulate notch ligands to support mammalian intestinal homeostasis

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Mouse fetal intestinal organoids: new model to study epithelial maturation from suckling to weaning

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LYVE-1

The atypical chemokine receptor 2 limits renal inflammation and fibrosis in murine progressive immune complex glomerulonephritis

Bideak A, et al. (2018). *Kidney Int.* DOI: 10.1016/j.kint.2017.11.013

High-dimensional cell-level analysis of tissues with Ce3D multiplex volume imaging

Li W, et al. (2019). *Nat Protoc.* DOI: 10.1038/s41596-019-0156-4

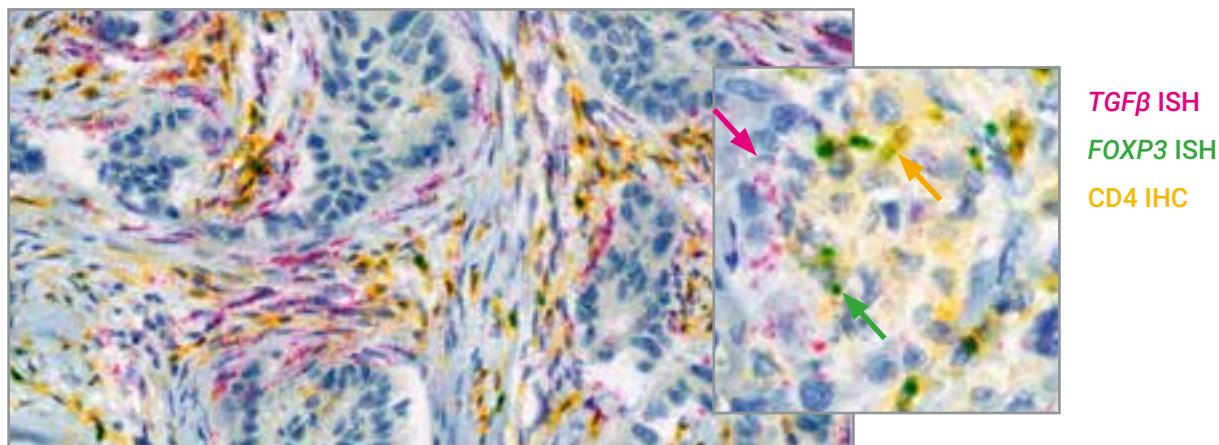


FIGURE 8. Detection of regulatory T cells in the TME of human lung cancer. The RNAscope VS Duplex Assay for *TGFB* and *FOXP3* was combined with IHC for CD4.

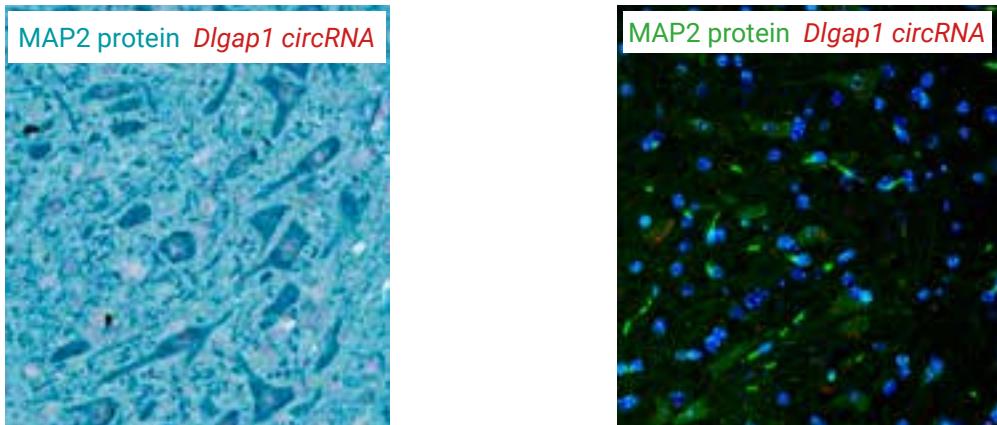


FIGURE 9. Cell type-specific expression of circRNAs in the tissue. The BaseScope™ assay for the circRNA *Dlgap1* was combined with IHC for Map2. Due to the fluorescent nature of the Fast Red chromogen, the image was visualized under brightfield (left) and fluorescent (right) microscopy.

MBP

Oligodendroglia are particularly vulnerable to oxidative damage after neurotrauma *in vivo*

Giacci MK, et al. (2018). *J Neurosci*. DOI: 10.1523/JNEUROSCI.1898-17.2018

NRG1 type I dependent autocrine stimulation of Schwann cells in onion bulbs of peripheral neuropathies

Fledrich R, et al. (2019). *Nat Commun*. DOI: 10.1038/s41467-019-09385-6

MCH

Control of feeding behavior by cerebral ventricular volume transmission of melanin-concentrating hormone

Noble EE, et al. (2018). *Cell Metab*. DOI: 10.1016/j.cmet.2018.05.001

Melanin-concentrating hormone neurons promote rapid eye movement sleep independent of glutamate release

Naganuma F, et al. (2019). *Brain Struct Funct*. DOI: 10.1007/s00429-018-1766-2

Functional analysis reveals differential effects of glutamate and MCH neuropeptide in MCH neurons

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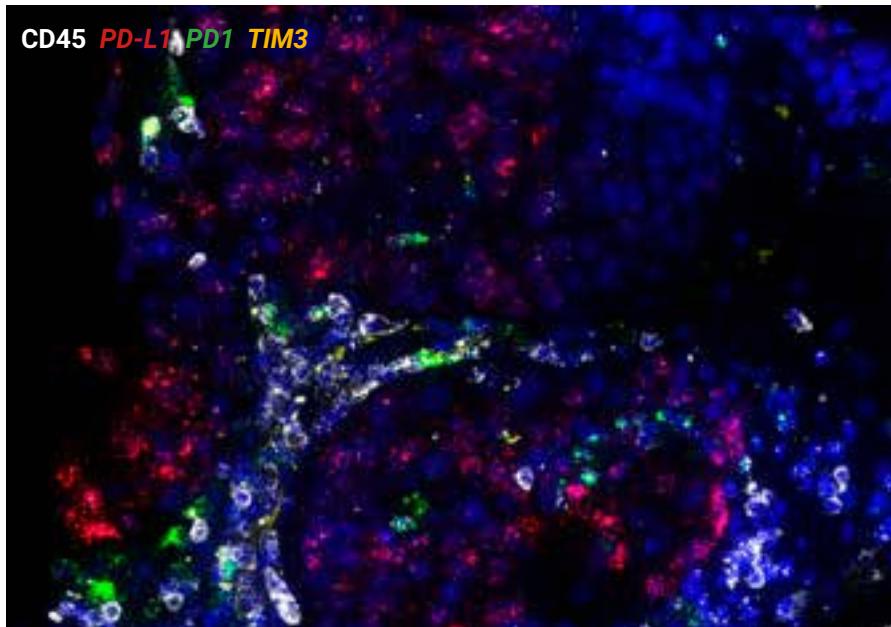


FIGURE 10. Detection of multiple immune checkpoint markers in the TME of human lung cancer. The RNAscope Multiplex Fluorescent V2 Assay for *PD-L1*, *PD1*, and *TIM3* was combined with IF for *CD45*.

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