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Reporting Summary

Nature Research wishes to improve the reproducibility of the work that we publish. This form provides structure for consistency and transparency in reporting. For further information on Nature Research policies, see our Editorial Policies and the Editorial Policy Checklist.

For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.

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n/a	Confirmed
	$oxed{\boxtimes}$ The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
	🔀 A statement on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly
	The statistical test(s) used AND whether they are one- or two-sided Only common tests should be described solely by name; describe more complex techniques in the Methods section.
\boxtimes	A description of all covariates tested
	🔀 A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
	A full description of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) AND variation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)
	For null hypothesis testing, the test statistic (e.g. <i>F</i> , <i>t</i> , <i>r</i>) with confidence intervals, effect sizes, degrees of freedom and <i>P</i> value noted <i>Give P values as exact values whenever suitable.</i>
\boxtimes	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
\boxtimes	Estimates of effect sizes (e.g. Cohen's <i>d</i> , Pearson's <i>r</i>), indicating how they were calculated
	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.
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Software and code

Policy information about <u>availability of computer code</u>

Data collection

BD FACSDiva™, Summit™, LAS X (Version 3.3.0.16799)

Data analysis

Cytoscape (Version 3.6.1), FlowJo X, R (Version 3.5.1), RStudio (Version 1.1.453), GraphPad Prism 7.0, Microsoft Office Professional Plus 2016. Transcriptomic data was performed in RStudio using the following packages: limma (Version 3.45.7), GSVA (Version 1.37.2), NicheNet, survminer (Version 0.4.7), Seurat (Version 3.2.2)

For manuscripts utilizing custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors and reviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Research guidelines for submitting code & software for further information.

Data

Policy information about availability of data

All manuscripts must include a data availability statement. This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A list of figures that have associated raw data
- A description of any restrictions on data availability

Transcriptomic data was downloaded from the NCBI GEO database (https://www.ncbi.nlm.nih.gov/geo/) with the following accession numbers: GSE1547117, GSE1651518, GSE6245219, GSE7172920. The single cell RNA-Seq data was downloaded from the Genome Sequence Archive (https://bigd.big.ac.cn/gsa/) under the project PRJCA00106358. The data from the PAAD project of the TCGA is available for download from the Broad Institute GDAC Firehose (https:// gdac.broadinstitut.org)28. The transcriptomic data published by Puleo et al was downloaded from the ArrayExpress database with the accession number E-MTAB-613426. Data from the PACA-AU project of the ICGC (release 25) was obtained from https://dcc.icgc.org/27. The remaining data are available within the article, supplementary information or available from the authors upon request.

Field-specific reporting
Please select the one below that is the best fit for

•	cinc reporting
	ne below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.
Life sciences	Behavioural & social sciences
TOT a reference copy of t	the document with an sections, see <u>nature.com/documents/m-reporting-summary-nat.pdr</u>
Life scier	nces study design
	close on these points even when the disclosure is negative.
Sample size	No statistical method was used to pre-determine sample size. In the experiments where human monocytes or macrophages were used, at
·	least cells derived from 4 healty donors was used: Co-Culture of macrophages with cell lines n=4, Dendrimer stimulation of macrophages n = 7.
Data exclusions	Data was excluded from analysis only if positive controls did not work properly, as it could be for the standard curve in a cytokine ELISAs.
Replication	Each experiment performed in this paper has been successfully repeated independently at least 3 times.
Randomization	No randomization was used.
Blinding	Researchers were not blinded when conducting analysis of samples.
	·
Antibodies	ChIP-seq
Eukaryotic	
	ogy and archaeology MRI-based neuroimaging d other organisms
	earch participants
Clinical dat	a
Dual use re	esearch of concern
Antibodies	
Antibodies used	AJoJo1
	##Flow Cytometry and Microscopy## Anti-PanCytokeratine - AlexaFluor 488
	Anti-CD163 - Brilliant Violet 421™ Anti-Alpha-Smooth Muscle Actin - eFluor® 570
	Anti-DC-SIGN (AZN-D1) - AlexaFluor 488
	Anti-HLA-DR - BV786 Anti-E-Cadherin - AlexaFluor 647
	Anti-Vimentin - AlexaFluor 488 Goat anti rabbit IgG - AlexaFluor 488
	Anti LewisY antibody
	Anti LewisX antibody Anti-mouse IgM - FITC
	Anti-mouse IgG - FITC Anti-human IgG - FITC
	##Western Blot##
	CA19-9 Antibody (SPM110)
	Purified anti-human E-Cadherin (24E10) Purified anti-human Vimentin
	Purified anti-human TMEM30B Purified anti-human MAL2
	Purified anti-human Rab25

Purified anti-human GALNT3 Antibody

Polyclonal anti-human ZEB1 Polyclonal anti-human ZEB2 Purified anti-β-actin Antibody Goat anti-rabbit - HRP Rabbit anti-mouse - HRP Goat anti-sheep - HRP

##ELISA##

Capture Antibody - IL-10
Detection Antibody - IL-10
Capture Antibody - IL-6
Detection Antibody - IL-6
Capture Antibody - TNFa
Detection Antibody - TNFa

Validation

Statements in antibodies data sheets provided by manufacturer. All antibodies were optimized and validated by serial dilutions.

Eukaryotic cell lines

Policy information about cell lines

Cell line source(s)

ASPC1, Mia PaCa-2 and PL45 were acquired from ATCC. BxPC3 are a kind gift from Dr. A. Frampton (Imperial College, London, UK). PaTuS and PaTuT are a kind gift from Dr. I. van Die (Amsterdam UMC, The Netherlands).

Authentication Cell lines were tested were tested for their authentication by STR-PCR, performed by BaseClear (Leiden, The Netherlands),

previous to the start of the project.

Mycoplasma contamination Cell lines were routinely tested for Mycoplasma using PCR, resulting all negative.

Commonly misidentified lines (See ICLAC register)

No Commonly misidentified cell lines were used in this paper.

Human research participants

Policy information about studies involving human research participants

Population characteristics

Describe the covariate-relevant population characteristics of the human research participants (e.g. age, gender, genotypic information, past and current diagnosis and treatment categories). If you filled out the behavioural & social sciences study

design questions and have nothing to add here, write "See above."

Recruitment

Biopsies were collected from patients that undergo to whipple surgery. Samples with a pathological assessment of PDAC

were used. For the analysis of sialic acids in normal and tumor tissue, samples containing adjacent normal tissue were

analyzed. No bias that could affect the results were identified.

Ethics oversight Medical Ethical Committee from the Amsterdam UMC, Location VUmc.

Note that full information on the approval of the study protocol must also be provided in the manuscript.

Flow Cytometry

Plots

Confirm that:

The axis labels state the marker and fluorochrome used (e.g. CD4-FITC).

The axis scales are clearly visible. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers).

All plots are contour plots with outliers or pseudocolor plots.

A numerical value for number of cells or percentage (with statistics) is provided.

Methodology

Sample preparation For the analysis of cell lines, cells were harvested using a mix of EDTA/Trypsin.

Instrument CyAn™ ADP Analyzer or BD LSRFortessa™.

Software Acquisition: BD FACSDiva™ for Fortessa, Summit™ Software for CyAn.

Analysis: FlowJo X.

Cell population abundance

Describe the abundance of the relevant cell populations within post-sort fractions, providing details on the purity of the samples and how it was determined.

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 For the determination of glycan structures in PDAC cell lines, only single live cells were analyzed by gating in Viability Dye negative cells.