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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.

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For all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or inlethods section.
n/a Confirmed
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.
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 Give P values as exact values whenever suitable.
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
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.
Software and code

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

Data collection

Provide a description of all commercial, open source and custom code used to collect the data in this study, specifying the version used OR state that no software was used.

Data analysis

Provide a description of all commercial, open source and custom code used to analyse the data in this study, specifying the version used OR state that no software was used.

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 <u>availability of data</u>

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

The accession numbers for the nucleotide and protein sequences for variable regions of T1AM09 and L1AM04 are GenBank: MZ600149 - MZ600152. Glycan array data were deposited in the NCBI Gene Expression Omnibus (GEO) database with accession number GSE180517.

Field-specific reporting				
<u>.</u>		s the best fit for your research. If you are not sure, read the appropriate sections before making your selection.		
✓ Life sciences		ehavioural & social sciences		
For a reference copy of t	the document with	all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>		
Life scier	nces stu	udy design		
All studies must dis	sclose on these	points even when the disclosure is negative.		
Sample size	We selected samples from our cohort with available PBMC and high anti-AM titers.			
Data exclusions	No data were ex	No data were excluded from the analysis		
Replication	All attempts of replication have been successful			
Randomization	NA NA			
Blinding	NA			
Reportin	g for sr	pecific materials, systems and methods		
		about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material,		
system or method list	ted is relevant to	your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.		
Materials & exp		<u> </u>		
n/a Involved in th	,	n/a Involved in the study		
Antibodies		∑ ChIP-seq		
Eukaryotic	ogy and archaeol	ogy MRI-based neuroimaging		
	nd other organism			
	search participant			
Clinical dat				
Dual use re	esearch of concer	n		
·				
Antibodies				
Antibodies used		nti-human IgG HRP 2048-05 I1911-RJ35G Southern Biotech		
		nti-human IgG HRP 2048-05 B0819-Q380B Southern Biotech anti-Human IgG 2040-03 K0211-MD587 Southern Biotech		
	Surelig	ht P3 Cy5 D7-2212 P3001-09-029 Columbia Biosciences		
		anti-human IgG (H+L), Mouse ads-UNLB 6145-01 Southern Biotech nti-human IgG-Biotin 209-065-098 118179 Jackson Immunoresearch		
		nti-human CD3 (B209500) Biolegend		
	PE/Cy7	anti-human CD19 (B242979) Biolegend		
		nti-human CD14 (B247208) Biolegend Iti-human IgG Fc (B232398) Biolegend		
Validation	All antibodies used in this study were verified for use for ELISA, Fluorescent Microscopy, or Flow Cytometry by the manufacturer. Secondary antibodies were titrated for optimal performance.			
	(1-1-1)			
Eukaryotic c	ell lines			
Policy information about <u>cell lines</u>				
		293T and F cell lines were used for protein expression and obtained as a generous gift from Dr. Kartik Chandran and Dr. Jonathan Lai at Albert Einstein College of Medicine.		

The cell lines used were authenticated

Authentication

Mycoplasma contamination

Cell lines were not tested for mycoplasma contaminaton

Commonly misidentified lines (See ICLAC register)

Name any commonly misidentified cell lines used in the study and provide a rationale for their use.

Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research

C57BL/6 female mice (7–8 weeks old; Charles River) Laboratory animals

Wild animals The study did not involve wild animals

The study did not involve field-collected samples Field-collected samples

Ethics oversight The animal studies were approved by the Institutional Animal Care and Use Committee at the Albert Einstein College of Medicine.

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

Human research participants

Policy information about studies involving human research participants

Both human subjects selected for this study were healthy adults with confirmed Mtb infection/exposure Population characteristics

These subjects were recruited as part of an ongoing TB immunology study. These subjects were selected because of available Recruitment

PBMC and high antibody titers to our antigen of interest.

Ethics oversight The study was approved by the institutional review board of the Albert Einstein College of Medicine. All human subjects gave written informed consent before enrollment.

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

Clinical data

Policy information about clinical studies

All manuscripts should comply with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.

Clinical trial registration NA; this is not a clinical trial

Einstein IRB protocol #2006-428; CSU IRB approval numbers 15-104B and 09-006B Study protocol

NA Data collection

Outcomes NA

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

Human PBMC isolated by Ficoll separation Sample preparation

Instrument Becton Dickinson FACSAria high-speed cell sorter

FlowJo was used to analyze the software Software

We detect about 1.5-2% antigen positive B cells and sort into individual wells, Cell population abundance

Lymphocytes were first identified (FSC/SSC), then memory B cells were further enriched based on CD20+/CD19+ and CD27+. Gating strategy

Secondary staining using streptavidin labeled with phycoerythrin (PE) was used to detect the AM-positive population.

Fluorescence-minus-one (FMO) controls were used to set the sorting gates for double positive (AM+, IgG+) B cells. More details are provided in the methods and supplemental figures.

Tick this box to confirm that a figure exemplifying the gating strategy is provided in the Supplementary Information.