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	Elias Haddad
Corresponding author(s):	Virginie Tardif
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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, seeAuthors & Referees and theEditorial Policy Checklist.

Statistics			
For all statistical analys	es, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.		
n/a Confirmed			
The exact sam	pple size $(n)$ for each experimental group/condition, given as a discrete number and unit of measurement		
A statement of	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	X A description of all covariates tested		
A description	of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons		
	ion of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)		
	hesis testing, the test statistic (e.g. $F$ , $t$ , $r$ ) with confidence intervals, effect sizes, degrees of freedom and $P$ value noted exact values whenever suitable.		
For Bayesian a	analysis, information on the choice of priors and Markov chain Monte Carlo settings		
For hierarchic	al and complex designs, identification of the appropriate level for tests and full reporting of outcomes		
Estimates of e	effect sizes (e.g. Cohen's $d$ , Pearson's $r$ ), indicating how they were calculated		
	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.		
Software and c	ode		
Policy information abou	ut availability of computer code		
Data collection	n/a		
Data analysis	n/a		
For manuscripts utilizing custo	om algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors/reviewers. deposition in a community repository (e.g. GitHub). See the Nature Research guidelines for submitting code & software for further information.		
Data			
Policy information abou	ut <u>availability of data</u>		
- Accession codes, un - A list of figures that	include a <u>data availability statement</u> . This statement should provide the following information, where applicable: ique identifiers, or web links for publicly available datasets have associated raw data restrictions on data availability		
Sequence and gene expre	ession data are available at the Gene Expression Omnibus (accession number pending).		
Field-speci	fic reporting		
Please select the one b	elow that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.		
<b>x</b> Life sciences	Behavioural & social sciences Ecological, evolutionary & environmental sciences		

For a reference copy of the document with all sections, see  $\underline{\mathsf{nature}.\mathsf{com}/\mathsf{documents}/\mathsf{nr}-\mathsf{reporting}-\mathsf{summary}-\mathsf{flat}.\mathsf{pdf}}$ 

### Life sciences study design

All studies must dis	sclose on these points even when the disclosure is negative.
Sample size	Sample sizes were not predetermined based on statistical methods, but were chosen according to the standards of the field (at least four independent human tonsil samples were used for each condition, most times 7-10 were used). This is due to previous work and takes into account the heterogeneity of human samples.
Data exclusions	no data was excluded from consideration.
Replication	Reported results were consistently replicated across multiple experiments with all replicates generating similar results. There were at least 2 replicates for each experiment, usually 4-6 replicates.
Randomization	Each human tonsil was used for all experimental subgroups so there were both autologous and heterologous controls for each human subject.
Blinding	When performing ELISA and Luminex experiments on co-culture supernatants, samples were blinded and given a number to keep from bias.

### Reporting for specific materials, systems and methods

Methods

We require information from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, system or method listed 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 & experimental	l systems Methods
n/a Involved in the study	n/a Involved in the study
Antibodies	<b>▼</b> ChIP-seq
<b>x</b> Eukaryotic cell lines	Flow cytometry
<b>x</b> Palaeontology	MRI-based neuroimaging
Animals and other organi	sms
Human research participa	ants
Clinical data	
·	
Antibodies	
Antibodies used	All antibodies are listed in supplementary information with supplier name, clone number and catalog number
Validation	antibodies were used as per manufacturers recommendation for concentration and titrated as necessary
Human research par	ticipants
Policy information about studies	s involving human research participants
	Tonsils were obtained from patients age 17-50 with a mix of both males and females. Samples used were from patients who were having their tonsils removed due to chronic infections.

Participants were recruited who were having their tonsils taken removed due to repetitive chronic infections of the area Recruitment

Martin Memorial Health Systems (Florida), St. Christopher's Hospital for Children (Pennsylvania) and National Institutes of Health Ethics oversight Clinical Research Center (Bethesda). The Institutional Review Boards at the relevant institutions approved all procedures, and all participants provided signed informed consent.

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 ICMJEguidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.

Clinical trial registration	ClinicalTrials.gov Identifier: NCT00001316
Study protocol	under the link above and using the identifier listed, one can access the full study protocol
Data collection	LN mononuclear cells collected from HIV+ patients for use in our study

Outcomes n/a

#### 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	Sample preparation listed in Methods
Instrument	FACS Aria Fusion, and Fortessa
Software	FACSDiva for collection and data analyzed using FlowJo (v10)
Cell population abundance	100-200 million TMNC were sorted per patient to acquire relevant cell populations: 300,000 GCB and GCTfh, 1,000 ILCFR, 15,000 Treg, 3,000 ILC3
	Purity was assessed by staining for cell specific markers and validated by RNAseq data
Gating strategy	Relevant gating strategies shown in Figure 1 and the Supplementary Information

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