Data S1: COMBAT Consortium Author Details and Contributions

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COMBAT Consortium author contributions

1. Patient recruitment and cohorts

Alexander J Mentzer, Mark Ainsworth, Alice Allcock, Carolina V Arancibia-Cárcamo, J Kenneth Baillie, Brian Angus, Archana Bashyal, Sally Beer, Tihana Bicanic, Andrew Brent, Andrew Brown, Christopher P Conlon, Neil Davidson, Julie Dequaire, Susanna J Dunachie, Alexis Espinosa, Rory Fairhead, Shayan Fassih, John Frater, Maria Fernandez Mendoza, Thomas Foord, Anastasia Fries, Veronica Gallardo Sanchez, Dominique Georgiou, Charles Hinds, Clare Hird, Paula Hutton, Katie Jeffery, David Kim, Paul Klenerman, Julian C Knight, Andrew Kwok, Teresa Lockett, Jose Martinez, Philippa C Matthews, Stuart McKechnie, Denise O'Donnell, Alex Novak, Ian Pavord, Elena Perez, Thomas Ritter, Matthew Rowland, Malcolm G Semple, Donal Skelly, Alberto Sobrinodiaz, Lizzie Stafford, Adan Taylor, Hannah Thraves, Lance Turtle, Holm Uhlig, Peter Watkinson, Rebecca K Young, Jonathan Youngs.

COVID-19 patients (Oxford)

Patient recruitment, consent and phenotyping by Alexander J Mentzer, Mark Ainsworth, Brian Angus, Archana Bashyal, Sally Beer, Neil Davidson, Julie Dequaire, Alexis Espinosa, Rory Fairhead, Shayan Fassih, Maria Fernandez Mendoza, Thomas Foord, Anastasia Fries, Veronica Gallardo Sanchez, Dominique Georgiou, Clare Hird, Paula Hutton, David Kim, Andrew Kwok, Teresa Lockett, Jose Martinez, Elena Perez, Thomas Ritter, Alberto Sobrinodiaz, Adan Taylor, Hannah Thraves, Peter Watkinson, Rebecca K Young. Patients were recruited as part of Sepsis Immunomics (Julian C Knight, Chief Investigator for Sepsis Immunomics; ISARIC/WHO Clinical Characterisation Protocol for Severe Emerging Infections (J Kenneth Baillie, Paul Klenerman, Malcolm G Semple, Lance Turtle); OPTIC study (Carolina V Arancibia-Cárcamo, Christopher P Conlon, Susanna J Dunachie, John Frater, Katie Jeffery, Philippa C Matthews, Denise O'Donnell, Donal Skelly, Lizzie Stafford).

Influenza patients and COVID-19 patients (non-Oxford)

Patients were recruited by Jonathan Youngs, members of the AspiFlu team and clinical research nurses. Supervised by Tihana Bicanic (Chief Investigator for AspiFlu).

Sepsis patients

Sepsis patients were recruited by Andrew Kwok, Sally Beer and the clinical research nurse team at Oxford University Hospitals Trust, with supervision and input by Julian C Knight (Chief Investigator for Sepsis Immunomics), Stuart McKechnie (local PI), Charles Hinds, Alexander J Mentzer, Andrew Brent, Alex Novak and Matthew Rowland.

Healthy volunteers

Healthy volunteers were recruited and sampled by Andrew Kwok with support from Alexander J Mentzer, Andrew Brown and Alice Allcock. Supervised by Julian C Knight as Chief Investigator (healthy volunteer study).

Supervision of recruitment and cohorts

Overall supervision and direction by Alexander J Mentzer, with input from Julian C Knight and Andrew Kwok.

2. Sample processing and extraction

Alexander J Mentzer, Alice Allcock, Chris Allan, Amy Beveridge, Sagida Bibi, Tihana Bicanic, Luke Blackwell, Andrew Brown, David Buck, Susana Camara, Elizabeth Clutterbuck, Fabiola Curion, Christina Dold, Tao Dong, Hal Drakesmith, Sally Felle, Leila Godfrey, Angie Green, Jennifer Hill, Elizabeth Jones, Julian C Knight, Andrew Kwok, Angela Lee, Aline Linder, Lorne Lonie, Maria Lopopolo, Spyridoula Marinou, Adam Mead,

Yanchun Peng, Andrew J Pollard, Laura Silva-Reyes, Christina Simoglou Karali, Christine S Rollier, Giuseppe Scozzafava, Hubert Slawinski, Lisa Stockdale, Amy Trebes, Marije Verheul, Lorna Witty Katherine Wray, Zixi Yin, Jonathan Youngs.

Sample processing

Preparation of samples, separation and cryopreservation of peripheral blood mononuclear cells Chris Allan, Amy Beveridge, Sagida Bibi, Luke Blackwell, Susana Camara, Elizabeth Clutterbuck, Christina Dold, Sally Felle, Angie Green, Jennifer Hill, Elizabeth Jones, Andrew Kwok, Aline Linder, Spyridoula Marinou, Yanchun Peng, Laura Silva-Reyes, Christina Simoglou Karali, Christine S Rollier, Marie Strickland, Lisa Stockdale, Marije Verheul, Katherine Wray, Zixi Yin, Jonathan Youngs.

Purification of genomic DNA and RNA

Alice Allcock, Andrew Brown, Andrew Kwok, Giuseppe Scozzafava.

Sample management, curation and cataloguing

Andrew Kwok, Alexander J Mentzer, Giuseppe Scozzafava.

Oxford Genomics Centre (OGC) core sequencing facility

Fabiola Curion, Angela Lee, Maria Lopopolo, Lorne Lonie, Hubert Slawinski, Amy Trebes, Lorna Witty

Supervision

Supervised in specific labs by Tihana Bicanic (London), David Buck (OGC), Tao Dong (WIMM), Julian C Knight (WHG), Andrew J Pollard (CCVTM); and with overall direction by Alexander J Mentzer and Julian C Knight.

3. Clinical phenotyping

Alexander J Mentzer, J Kenneth Baillie, Eleanor Barnes, Tihana Bicanic, Calliope A Dendrou, Lucy C Garner, Ling-Pei Ho, Julian C Knight, Andrew Kwok, Luke Jostins, T Phuong Quan, Fabian Ruehle, Stephen N Sansom, Malcolm G Semple, Alberto Santos Delgado, Lance Turtle, Jonathan Youngs.

4. Mass and flow cytometry

Giorgio Napolitani, Mariolina Salio, Claudia Monaco, Irina Udalova, David J Ahem, Yasemin- Xiomara Zurke, Zhichao Ai, Lea Dib, Rachel Etherington, Julian C Knight, Prathiba Kurupati, Andrew Kwok, Michalina Mazurczyk, Graham Ogg, Inhye Park, Mariana Periera Pinho, Emmanouela Repapi, Dominik Trzupek, Erinke van Grinsven, Lihui Wang.

Mass Cytometry - Whole Blood (figs 1, 3, 4, 5)

Experimental design and data generation

Andrew Kwok developed the antibody panel with help from Rachel Etherington and Michalina Mazurczyk and input from Giorgio Napolitani. Andrew Kwok processed the samples and performed the staining with help from Giorgio Napolitani. Rachel Etherington and Giorgio Napolitani acquired the CyTOF samples on the Helios Mass Cytometer.

Data analysis

Giorgio Napolitani performed manual de-barcoding and initial processing. Emmanouela Repapi performed processing and normalization of the data. Mariana Pereira Pino, and Emmanouela Repapi performed overall analysis with inputs from Giorgio Napolitani and Andrew Kwok. Mariana Pereira Pino performed cluster annotation and manual gating

analysis with inputs from Giorgio Napolitani. Emmanouela Repapi performed the statistical analysis.

Data integration:

Emmanouela Repapi performed the CITE-seq Mass Cytometry data integration.

<u>Supervision</u>

Giorgio Napolitani supervised the experimental design, data generation and data analysis with inputs from Julian C Knight.

Mass Cytometry - Myeloid panel (fig 2)

Experimental design and data generation

David J Ahern developed the myeloid CyTOF panel with input from Claudia Monaco, Irina Udalova, Lea Dib, Nikolaos Sousos and Lihui Wang. David J Ahern, Lea Dib and Inhye Park performed the staining.

Data analysis

Yasemin-Xiomara Zurke and David J Ahern performed the CyTOF analysis and clustering.

Supervision

Claudia Monaco and Irina Udalova supervised the data analysis and performed the interpretation of the data.

Multicolor Flow Cytometry

Data generation

Mariolina Salio developed and tested the flow cytometry panels. Mariolina Salio processed, stained and acquired the samples for Multicolor Flow Cytometry and did the data acquisition with help from Giorgio Napolitani.

Flow Cytometry Data Analysis

Mariana Pereira Pinho and Mariolina Salio performed data analysis with inputs from Giorgio Napolitani.

Supervision

Mariolina Salio supervised the experimental design, data generation and data analysis.

5. Whole blood total RNA-seq

Alexander J Mentzer, Moustafa Attar, Katie L Burnham, Emma E Davenport, James Docker, Clementine Geeves, Charles Hinds, Julian C Knight, Andrew Kwok, Angela Lee, Daniel O'Connor, Santiago Revale, Justin Whalley.

Data generation

Data was generated by Andrew Kwok, Moustafa Attar, James Docker, Clementine Geeves.

Data analysis

Data analysis was performed by Katie L Burnham, Andrew Kwok, Daniel O'Connor (WGCNA), Santiago Revale, Justin Whalley.

Supervision

Supervised by Alexander J Mentzer and Julian C Knight with Emma E Davenport and Charles Hinds.

6. 10X CITE-seq and repertoire

Stephen N Sansom, Calliope A Dendrou, Luke Jostins, Julian C Knight, Rachael Bashford-Rogers, Benjamin Fairfax, Dominik Aschenbrenner, Moustafa Attar, Paul Bowness, Adam P Cribbs, Fabiola Curion, Tao Dong, Ricardo Ferreira, Lucy C Garner, Maria Gomez Vazquez, Anna James-Bott, Ashwin Jainarayanan, Kathrin Jansen, Paul Klenerman, Piyush Kumar Sharma, Andrew Kwok, Angela Lee, Adam Mead, Alexander J Mentzer, Ruddy Montadon, Giorgio Napolitani, Isar Nassiri, Lauren Overend, Yanchun Peng, Frank Penkava, Bethan Psaila, Emmanouela Repapi, Santiago Revale, Jean-Baptiste Richard, Charlotte Rich-Griffin, Hubert Slawinski, Bo Sun, Chelsea Taylor, Supat Thongjuea, Orion Tong, Felicia A Tucci, Alexandru Voda, Guanlin Wang, Robert Watson, Hing Yuen Yeung, Yasemin-Xiomara Zurke.

Experimental design and data generation

Fabiola Curion, Julian C Knight, Andrew Kwok, Luke Jostins, Alexander J Mentzer, Rachael Bashford-Rogers and Stephen N Sansom contributed to sample selection and experimental design. Sample preparation, cell QC and single-cell capture was performed by Andrew Kwok, Hubert Slawinski and Moustafa Attar.

Analysis of the gene expression (GEX) data

Quantitation and genetic demultiplexing was performed by Fabiola Curion and Santiago Revale. Kathrin Jansen computed velocity matrices. Pre-processing and QC assessment of the GEX data was performed by Bo Sun, Fabiola Curion, Kathrin Jansen and Maria Gomez Vasquez with input from and Lauren Overend. Databasing of cell metrics was performed by Adam P Cribbs with help from Stephen N Sansom. Extraction of matrices for downstream analysis was performed by Stephen N Sansom. Alignment of the data was performed by Kathrin Jansen with advice from Supat Thongjuea and Stephen N Sansom. Clustering, marker and pathway analysis was performed by Stephen N Sansom. Initial clustering results were independently confirmed by Charlotte Rich-Griffin and Calliope A Dendrou. Analysis of viral transcripts was performed by Adam P Cribbs, Anna James-Bott and Lauren Overend.

Analysis of the antibody-derived tag (ADT) data

Analysis of the ADT data was performed by Fabiola Curion and Giorgio Napolitani with help from Emmanouela Repapi and input from Calliope A Dendrou and Stephen N Sansom.

Analysis of the CITE-seq and bulk BCR and TCR data

Santiago Revale performed CITE-seq VDJ pre-processing. Bo Sun performed single cell BCR repertoire analysis. Single cell TCR repertoire analysis with exploration of clonal size, integration with gene expression and Kmer usage was performed by Rob Watson, Orion Tong, Chelsea Taylor and Piyush Kumar Sharma. Felicia A Tucci performed the bulk BCR and TCR repertoire amplification, library preparation and sample handling. Rachael Bashford- Rogers performed the bulk BCR and TCR repertoire analysis. Repertoire team members who have provided guidance and feedback to the analyses and manuscript were Lauren Overend, Tao Dong, Charlotte Rich-Griffin, Yanchun Peng, Isar Nassiri, Paul Bowness, and Frank Penkava.

Multimodal annotation of the CITE-seq dataset

Annotation of B-cell subpopulations was performed by Bo Sun, Calliope A Dendrou and

Charlotte Rich-Griffin with input from Felicia A Tucci, Hing Yuen Yeung and Rachael Bashford-Rogers. Annotation of T/NK subpopulations was performed by Calliope A Dendrou, Lucy C Garner, Ricardo Ferreira, Giorgio Napolitani and Stephen N Sansom with input from Paul Klenerman, Andrew Kwok and Hing Yuen Yeung. Annotation of the myeloid subsets was performed by Andrew Kwok and Stephen N Sansom. Annotation of HSC and platelets was performed by Stephen N Sansom and Calliope A Dendrou with input from Bethan Psaila. Annotation of doublet populations was performed by Fabiola Curion and Kathrin Jansen. Initial marker visualization was performed by Charlotte Rich-Griffin and Fabiola Curion. Curation of annotations was performed by Stephen N Sansom with review from Andrew Kwok, Lucy C Garner and Calliope A Dendrou. Manuscript figures were prepared by Stephen N Sansom.

Composition analysis

Composition PCA and association testing was performed by Lucy C Garner and Charlotte Rich- Griffin with input from Calliope A Dendrou. Comparison of multiple differential abundance analysis methods and final analyses were performed by Charlotte Rich-Griffin and Lucy C Garner with input from Calliope A Dendrou. Clinical variable selection and covariate analysis were performed by Lucy C Garner, Charlotte Rich-Griffin and Calliope A Dendrou. Calliope A Dendrou interpreted the results and prepared final manuscript figures with input from Charlotte Rich-Griffin, Julian C Knight and Lucy C Garner.

Pseudobulk analysis

Initial pseudobulk counts were generated by Stephen N Sansom and assessed by Fabiola Curion and Luke Jostins. Normalized pseudobulk counts, principal components and UMAP co-ordinates were generated by Luke Jostins with advice from Stephen N Sansom and Fabiola Curion. Alexandru Voda performed differential expression analyses with input from Luke Jostins and Stephen N Sansom. Luke Jostins collected pathway gene lists with input from Lucy C Garner, Adam P Cribbs, Andrew Kwok, Jean-Baptiste Richard and Dominik Aschenbrenner, and carried out gene set enrichment analyses. Clinical covariate analysis was carried out by Luke Jostins with input from Lucy C Garner, Charlotte Rich-Griffin and Calliope A Dendrou. Luke Jostins interpreted results and generated manuscript figures with input from Stephen N Sansom, Calliope A Dendrou and JulianC Knight.

WGCNA and network analysis

Lucy C Garner performed initial exploratory analyses. WGCNA network analysis was performed by Charlotte Rich-Griffin, Fabiola Curion and Stephen N Sansom. Charlotte Rich-Griffin and Fabiola Curion identified sets of modules common to multiple cell types and characterized their relationship with the type I IFN signaling response. Stephen N Sansom tested module associations, performed additional analyses and interpreted the results with input from Calliope A Dendrou, Charlotte Rich-Griffin, Fabiola Curion, Andrew Kwok and Julian C Knight. Charlotte Rich-Griffin, Fabiola Curion and Stephen N Sansom prepared the figures.

Cell-cell interaction analysis

Cell-cell interaction was performed by Guanlin Wang with input from Supat Thongjuea.

Supervision

Luke Jostins, Julian C Knight, Rachael Bashford-Rogers and Stephen N Sansom supervised the experimental design. Rachael Bashford-Rogers supervised the BCR and repertoire analysis. Benjamin Fairfax supervised the TCR analysis. Luke Jostins supervised

the pseudobulk analysis and provided expert statistical guidance. Calliope A Dendrou supervised the composition analysis and co-supervised the multimodal annotation. Supat Thongjuea co- supervised the alignment work. Supat Thongjuea and Adam Mead supervised the cell-cell interaction analysis. Stephen N Sansom supervised the GEX preprocessing, alignment and clustering, the multimodal annotation and the WGCNA network analysis. Overall supervision and facilitation of work by Stephen N Sansom with Calliope A Dendrou (CITE-seq) and Rachael Bashford-Rogers (repertoire).

7. ATAC-seq

Tatjana Sauka-Spengler, Ivan Candido Ferreira, Martyna Lukoseviciute, Moustafa Attar, Santiago Revale, Julian C Knight, Andrew Kwok, Angela Lee, Hubert Slawinski, Michael Weinberger.

Data generation

Data was generated by Andrew Kwok, Moustafa Attar and Hubert Slawinski.

Data analysis

Data analysis was performed by Ivan Candido Ferreira, Martyna Lukoseviciute, Santiago Revale, Andrew Kwok, Tatjana Sauka-Spengler and Michael Weinberger.

Supervision

Tatjana Sauka-Spengler and Julian C Knight.

8. Genetics

Luke Jostins, Amanda Chong, Calliope A Dendrou, Damien Downes, Chris Eijsbouts, Ben Fairfax, Jim Hughes, Julian C Knight, Alexander J Mentzer, Yuxin Mi, Isar Nassiri, Stephen N Sansom, Ron Schwessinger, Ping Zhang.

Alexander J Mentzer and Andrew Kwok prepared samples for genotyping, Ben Hollis and Luke Jostins carried out quality control, principal component analysis and imputation. Amanda Chong carried out association analysis. Damien Downes, Chris Eijsbouts, Luke Jostins, Yuxin Mi, Isar Nassiri, Ron Schwessinger and Ping Zhang carried out integration with other data modalities. Calliope A Dendrou, Ben Fairfax, Jim Hughes, Luke Jostins, Julian C Knight, Alexander J Mentzer and Stephen N Sansom supervised specific analyses, and Luke Jostins provided overall supervision.

9. Proteomics (mass spectrometry)

Roman Fischer, Alberto Santos Delgado, Georgina Berridge, Philip Charles, Simon Davis, Raphael Heilig, Svenja Hester, Julian C Knight, Yuxin Mi, Darragh P O'Brien, Iolanda Vendrell.

Experimental design

lolanda Vendrell, Georgina Berridge and Roman Fischer developed the experimental design with inputs from Philip Charles, Simon Davis, Raphael Heilig.

Data generation

Georgina Berridge, Iolanda Vendrell, Svenja Hester, Darragh P O'Brien and Roman Fischer processed samples. Data was generated by Georgina Berridge, Iolanda Vendrell, Simon Davis and Raphael Heilig.

Data analysis

Roman Fischer and Philip Charles performed identification and quantitation. Yuxin Mi and Alberto Santos Delgado performed downstream analysis and interpretation.

Supervision

Iolanda Vendrell supervised the sample processing, Roman Fischer supervised the team.

Acknowledgements

The Proteomics team is grateful for the help of Zuzana Bencokova, Darren Blase and Benedikt Kessler (all TDI) for providing the environment for safe sample handling and analysis. We also thank Alexey Nesvizhskii and Fengchao Yu (both Department of Pathology. University of Michigan Medical School) for providing help and advice for data processing in Fragpipe.

10. Proteomics (Luminex)

Luzheng Xue, Wentao Chen, Yi-Ling Chen, David A Duncan, Paul Klenerman, Jian Luo, Claudia Monaco, Graham Ogg, Ian Pavord, Irina Udalova.

Experimental design

Wentao Chen and Luzheng Xue designed the Luminex assay panels with inputs from Claudia Monaco, Irina Udalova, Ian Pavord, Graham Ogg and Paul Klenerman.

Data generation

Yi-Ling Chen and Jian Luo processed the samples, performed the assays and generated the data.

Data analysis

Yi-Ling Chen, Jian Luo, Wentao Chen and David A Duncan performed data analysis.

Supervision

Luzheng Xue supervised the experimental design, data generation and data analysis.

11. Data integration

Julian C Knight, M Azim Ansari, Rachael Bashford-Rogers, Katie L Burnham, Helen Byrne, Mark Coles, Fabiola Curion, Calliope A Dendrou, Aden Forrow, Roman Fischer, Tanya Golubchik, Heather A Harrington, Ling-Pei Ho, Renee Hoekzema, Jim Hughes, Matthew A Jackson-Wood, Luke Jostins, Simon McGowan, Alexander J Mentzer, Yuxin Mi, Giorgio Napolitani, Emmanouela Repapi, Fabian Ruehle, Raphael Sanches Peres, Alberto Santos Delgado, Stephen N Sansom, Martin Sergeant, Anna Seigal, David Sims, Otto Sumray, Stephen Taylor, Adriana Tomic, Justin Whalley.

Clinical phenotyping

Statistical and machine learning analysis by Fabian Ruehle with input and interpretation from Ling-Pei Ho, Julian C Knight, Alexander J Mentzer and Stephen N Sansom.

<u>Integrative analysis of mass cytometery and CITEseq cell composition</u>

Analysis by Emmanouela Repapi and Yasemin-Xiomara Zurke with input from Fabiola Curion, Calliope A Dendrou, Giorgio Napolitani and Stephen N Sansom.

Similarity network fusion and patient subphenotyping

Analysis by Alberto Santos Delgado with input and interpretation from Katie L Burnham, Roman Fischer, Yuxin Mi, Julian C Knight.

Machine learning

Fabian Ruehle and Matthew A Jackson-Wood undertook machine learning analysis of phenotyping and experimental datasets with input from Ling-Pei Ho, Stephen N Sansom, Luke Jostins and Julian C Knight. Adriana Tomic performed analysis using Sequential Iterative Modeling "Over Night" (SIMON).

Topological data analysis

Otto Sumray and Renee Hoekzema performed the analysis with supervision by Heather A Harrington and input from Ling-Pei Ho and Stephen N Sansom.

Tensor and matrix decomposition

Data was analysed and interpreted by Justin Whalley, Heather A Harrington, Julian C Knight, Anna Seigal and Stephen N Sansom.

Data mining

Datasets identified and curated by Calliope A Dendrou with input from Guanlin Wang and Charlotte Rich-Griffin.

Data visualization

Martin Sergeant, Simon McGowan and Stephen Taylor developed software for data visualization and hosting through Multi Locus View (MLV), with input from Jim Hughes and Stephen N Sansom. Hallmark collation Katie L Burnham, Julian C Knight and Yuxin Mi with graphical visualization (Julian C Knight) and interactive visualization in MLV (Martin Sergeant, Stephen Taylor) with input from Stephen N Sansom. COMBAT logo and manuscript artwork Julian C Knight.

Supervision

Julian C Knight led and facilitated work by the group; Stephen N Sansom helped supervise the work.

12. Data management

Brian Marsden, John Broxholme, Adam P Cribbs, Robert Esnouf, Hai Fang, Hong Harper, Luke Jostins, Julian C Knight, Georgina Kerr, Paul Klenerman, Vinod Kumar, Alexander J Mentzer, Fiona Powrie, Stephen N Sansom, David Sims, John A Todd, Dapeng Wang.

Data Management

Hai Fang and Dapeng Wang managed data set registration and curation. Brian Marsden, Vinod Kumar and Hong Harper developed and implemented data management solutions. Alexander J Mentzer, Brian Marsden, Julian C Knight, Vinod Kumar, Georgina Kerr and Giuseppe Scozzafava implemented and managed processes for sample tracking and management. Brian Marsden and Luke Jostins designed the protocol for capture of project dataset metadata.

Digital Communications

Brian Marsden, Julian C Knight, Georgina Kerr, Stephen N Sansom and Dapeng Wang set up and managed communication platforms, website and channels for the consortium.

High-performance computing environment and Code management

A bespoke high-performance computational platform was provided by the University of Oxford's Biomedical Research Computing facility under the guidance of Robert Esnouf. Stephen N Sansom, Brian Marsden and Hai Fang helped configure and manage use of the platform with input from Adam P Cribbs. Stephen N Sansom documented use of the platform, configured the COMBATOxford Team Github and defined coding practice and management policies with input from Adam P Cribbs and David Sims.

Data management operational group

Brian Marsden, Hai Fang, Hong Harper, Julian C Knight, Vinod Kumar, Alexander J Mentzer, Dapeng Wang.

Data Access Committee

Julian C Knight, Paul Klenerman, Brian Marsden, Alexander J Mentzer, Fiona Powrie, John A Todd.

Supervision

Brian Marsden supervised the overall data management strategy and implementation with Julian C Knight. Brian Marsden, Stephen N Sansom and Georgina Kerr supervised development and use of the computational platform. Stephen N Sansom and Adam P Cribbs supervised management of code. Alexander J Mentzer supervised sample tracking and management.

13. Oversight and writing

Julian C Knight, Rachael Bashford-Rogers, Katie L Burnham, Richard Cornall, Fabiola Curion, Calliope A Dendrou, Benjamin Fairfax, Hai Fang, Roman Fischer, Lucy C Garner, Heather A Harrington, Charles Hinds, Luke Jostins, Paul Klenerman, Andrew Kwok, Brian Marsden, Helen McShane, Alexander J Mentzer, Yuxin Mi, Giorgio Napolitani, Graham Ogg, Andrew J Pollard, Fiona Powrie, Charlotte Rich-Griffin, Fabian Ruehle, Stephen N Sansom, Alberto Santos Delgado, Tatjana Sauka-Spengler, Gavin Screaton, Anna Seigal, Martin Sergeant, Stephen Taylor, Bo Sun, Felicia A Tucci, John A Todd, Adriana Tomic, Justin Whalley, Luzheng Xue.

Writing group

Julian C Knight, Rachael Bashford-Rogers, Calliope A Dendrou, Ben Fairfax, Roman Fischer, Luke Jostins, Paul Klenerman, Alexander J Mentzer, Giorgio Napolitani, Stephen N Sansom, Tatjana Sauka-Spengler, John A Todd, Luzheng Xue.

Living reports contributing to manuscript

Living reports on individual assay modalities writing leads Rachael Bashford-Rogers and Ben Fairfax (B and T cell repertoire); Katie L Burnham and Alexander J Mentzer (whole blood total RNA-seq); Fabiola Curion, Charlotte Rich-Griffin, Stephen N Sansom (WGCNA scRNA-seq); Calliope A Dendrou (10X CITE-seq composition), Roman Fischer (proteomics mass spectrometry); Heather A Harrington (topological data analysis); Luke Jostins (genetics); Alexander J Mentzer (clinical phenotyping); Giorgio Napolitani (flow and mass cytometry); Fabian Ruehle (statistical analysis clinical data, machine learning); Martin Sergeant and Stephen Taylor (Multi Experiment Viewer); Stephen N Sansom (10X CITE-seq pre-processing, alignment and clustering); Adriana Tomic (Sequential Iterative Modeling "Over Night" SIMON); Justin Whalley and Stephen N Sansom (Tensor analysis); Luzheng Xue (proteomics luminex). Contributions to reports by team members detailed within teams sections.

Manuscript writing, review and editing

Original draft written by Julian C Knight. Review and editing by Rachael Bashford-Rogers, Katie Burnham, Richard Cornall, Calliope A Dendrou, Hai Fang, Ben Fairfax, Roman Fischer, Lucy C Garner, Heather A Harrington, Charles Hinds, Luke Jostins, Andrew Kwok, Paul Klenerman, Alexander J Mentzer, Yuxin Mi, Giorgio Napolitani, Fiona Powrie, Stephen N Sansom, Alberto Santos Delgado, Anna Seigal, Bo Sun, Felicia A Tucci, John A Todd, Luzheng Xue. All other Consortium authors saw, had the opportunity to comment on, and approved the final draft.

Leadership and oversight

Julian C Knight led the COMBAT consortium including the work described in this paper. Paul Klenerman led the wider Oxford COVID-19 Immunology Research programme. Stephen N Sansom helped lead data analysis and interpretation. COMBAT team leadership and oversight Rachael Bashford-Rogers, Calliope A Dendrou, Ben Fairfax, Roman Fischer, Luke Jostins, Julian C Knight, Paul Klenerman, Brian Marsden, Alexander J Mentzer, Giorgio Napolitani, Stephen N Sansom, John A Todd, Tatjana Sauka-Spengler, Luzheng Xue. Georgina Kerr provided project management.

Steering committee

Richard Cornall, Fiona Powrie, Julian C Knight, Paul Klenerman, Alexander J Mentzer, Helen McShane, Graham Ogg, Andrew J Pollard, Gavin Screaton, John A Todd.

Consortium team membership, contributions (CRediT taxonomy), ORCID, funding, conflict of interest and acknowledgements

NAME	MEMBERSHIP OF COMBAT TEAM(S) (1) 10X CITE-seq; (2) ATAC-seq; (3) Whole blood total RNA-seq; (4) Clinical phenotyping; (5) Data integration; (6) Data management; (7) Flow and mass cytometry; (8) Genetics; (9) Serology and biochemistry; (10) Proteomics (Luminex); (11) Proteomics (mass spectrometry); (12) Repertoire; (13) Sample processing; (14) Viral sequencing; (15) Recruitment and sampling; (16) Oversight and strategic	CONTRIBUTION (CRediT taxonomy) ¹ (1) Conceptualization; (2) Methodology; (3) Software; (4) Validation; (5) Formal analysis; (6) Investigation; (7) Resources; (8) Data curation; (9) Writing-original draft; (10) Writing-review and editing; (11) Visualization; (12) Supervision; (13) Project administration; (14) Funding acquisition.	ORCID iD	FUNDING	CONFLICT OF INTEREST	ACKNOWLEDGEMENTS
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Alice Allcock (A.A.)	13,15	6,7		-		
Brian Angus (B.A.)	13	7				
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Georgina Kerr (G.K.)	6,16	13			
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Julian C Knight (J.C.K.)	1,3,4,5,6,8,11,13,15,16	1,2,4,5,6,7,8,9,10,11,12,1 3,14	0000-0002-0377-5536	The research was supported by the COVID-19 Research Response Fund, University of Oxford, Wellcome Trust Core Award Grant Number 203141/Z/16/Z and the NIHR Oxford Biomedical Research Centre. The views expressed are those of the author(s) and not necessarily those of the NHS, the NIHR or the Department of Health. JCK is supported by Wellcome	Wellcome Centre for Human Genetics Flow Cytometry Facility
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Angela Lee (A.L.)	1,2,3,14	2,4,12,13,14	0000-0002-7591-825X	Wellcome Trust [Core Award 203141/Z/16/Z]	Gene Well Hum by W refer for th initia	hank the Oxford comics Centre at the come Centre for an Genetics (funded /ellcome Trust grant ence 203141/Z/16/Z) ne generation and I processing of encing data
Aline Linder (A.Li)	13	6			3343	J

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Lorne Lonie (L.L.)	13	12		Wellcome Trust [Core Award 203141/Z/16/Z]	
Maria Lopopolo (M.L.)	13	6,7	0000-0001-8502-9264	Wellcome Trust [Core Award 203141/Z/16/Z]	
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Denise O'Donnell (D.O'D.)	15	7		
Graham Ogg (G.O.)	7,10	1		
Lauren Overend (L.O.)	1,12	1,2,3,6	0000-0002-2802-048X	Wellcome Trust
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lan Pavord (I.Pav.)	10	1		NIHR Oxford BRC

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Mariana Pereira Pinho (M.P.P.)	7	2,4,5,8,10,11	0000-0002-5759-2436	MRC HIU core funding		
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Fiona Powrie (F.Po.)	6,16	12				
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Jean-Baptiste Richard (J-B.R.)	1,5	5	0000-0002-4052-9772	
Charlotte Rich- Griffin (C.R-G.)	1	2,3,5,6,7,9,11	0000-0001-8212-9542	NIHR Oxford BRC (BRCRCF19-04)
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Christine S Rollier (C.S.R.)	13	6	0000-0002-9712-8080	NIHR Oxford BRC		
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Mariolina Salio (M.S.)	7	1,2,4,5,6,9,10,11,12	0000-0002-4821-3801	MRC HIU core funding		
Stephen Nicholas Sansom (S.N.S.)	1,4,5,6,16	1,2,3,4,5,6,7,8,9,10,11,12,	0000-0003-4569-8513	Kennedy Trust for Rheumatology Research, KENN171803 Core support		
Raphael Sanches Peres (R.S.P.)	1	5				
Alberto Santos Delgado (A.S.D.)	4,5,11	1,2,3,4,5,6,12	0000-0002-9163-7730			
Tatjana Sauka- Spengler (T.S-S.)	2,16	2,5,12				
Ron Schwessinger (R.S.)	8	5				
Giuseppe Scozzafava (G.S.)	13	6,7		RRF BRD00230		
Gavin Screaton (G.Sc.)	9	12		WT SIA and Schmidt Futures	Sits on GSK Vaccines SAB, founder RQ Biotechnolo gy	
Anna Seigal (A.S.)	5	1,2			3)	

Martin Sergeant	5,6	1,6,7,8,12,13,14	0000-0001-9700-0418	Medical Research	MGS reports grants from DHSC, NIHR UK, MRC UK, HPRU in Emerging and Zoonotic Infections, and University of Liverpool, during the conduct of the study; and is chair of the Infectious Diseases Science Advisory Board and minority shareholder of Integrum Scientific, Greensboro NC, outside the submitted work.	
(M.S.)				Research Council (MRC) Core Funding (MC_UU_00016 /14), National		

				Institutes of Health (USA) grant number	
				R24DK106766	
Christina Simoglou Karali (C.S.K.)	1,13	6,7	0000-0003-2169-6580		
David Sims (D.S.)	1	12		204826/Z/16/Z	
Donal Skelly (D.Sk.)	15	7	0000-0002-2426-3097	DS is an NIHR Academic Clinical Fellow	
Hubert Slawinski (H.S.)	1,2,13	6,7	0000-0002-8996-8477	Wellcome Trust [Core Award 203141/Z/16/Z]	
Alberto Sobrinodiaz (A.S.)	15	6,7		NIHR Clinical Research Network	
Nikolaos Sousos (N.S.)	1,13	6,7	0000-0002-9100-2954	CRUK Clinical Research Training Fellowship (C130623/A249 471)	
Lizzie Stafford (L.S.)	15	7			
Lisa Stockdale (L.St.)	13	6	0000-0002-2576-8783		
Marie Strickland (M.St.)	1,13	6,7			
Otto Sumray (O.S.)	5	2,3,5	0000-0003-1735-0595	UK Centre for Topological Data Analysis EPSRC grant EP/R018472/1	
Bo Sun (B.S.)	1,12	1,2,3,5,6,8,9,10,11	0000-0002-5507-6657	Patrick Berthoud Charitable Trust	
Chelsea Taylor (C.T.)	12	1,2,3,5,6,8,10,11	0000-0002-1635-4167		

Stephen Taylor (S.T.)	5	1,11,12	0000-0002-3559-4334	Medical Research Council (MRC) Core Funding (MC_UU_00016 /14)		
Adan Taylor (A.T.)	15	6,7		,		
Supat Thongjuea (S.Th.)	1	12	0000-0002-9129-4694			
Hannah Thraves (H.T.)	15	6,7		NIHR Clinical Research Network		
John A Todd (J.A.T.)	6,16	12,14	0000-0003-2740-8148	Wellcome Core Award Grant Number 203141/Z/16/Z supporting the Oxford Genomics Centre in the Wellcome Centre for Human Genetics, University of Oxford, and a Strategic Award to the Diabetes and Inflammation Laboratory from the JDRF (4- SRA-2017-473- A-A) and the Wellcome (107212/A/15/Z)	JAT is a member of the Human Genetics Advisory Board of GSK	
Adriana Tomic (A.T.)	5	2,3,5,6,8,12	0000-0001-9885-3535			
Orion Tong (O.T.)	12	2,3,5,6,9,11	0000-0002-0659-5944			

Amy Trebes (A.Tr.)	14	6,7	0000-0002-3715-9109	Wellcome Trust	
				[Core Award 203141/Z/16/Z]	
Dominik Trzupek (D.T.)	7	7			
Felicia Anna Tucci (F.A.T.)	1,12	1,2,9,11,12	0000-0001-5813-9708	Wellcome Trust	
Lance Turtle (L.T.)	4,6,15	6,7,8,12,13,14	0000-0002-0778-1693	NIHR HPRU EZI, ISARIC4C MRC, Wellcome fellowship 205228/Z/16/Z	
Irina Udalova (I.U.)	7	1,5,6,10,12,13,14			
Holm Uhlig (H.U.)	15	7			
Erinke van Grinsven (E.V.G.)	7	6			
Iolanda Vendrell (I.V.)	11	2,4,6,10,12	0000-0001-7187-4061	COI	
Marije Verheul (M.V.)	13	6			
Alexandru Voda (A.V.)	1	3,5	0000-0003-2974-6992		
Guanlin Wang (G.W.)	1	5	0000-0002-5622-3890		
Lihui Wang (L.W.)	7	6			
Dapeng Wang (D.W.)	6	8	0000-0002-9925-4574		
Peter Watkinson (P.W.)	15	7			
Robert Watson (R.W.)	12	1,2,3,5,6,8,10,11	0000-0002-8838-9406		
Michael Weinberger (M.W.)	1,2	5	0000-0001-8235-6825	BHF Programme Grant RG/13/9/303269	
Justin Whalley (J.W.)	3,5,14	2,3,5	0000-0002-0403-3091		

Lorna Witty (L.W.)	13	12		Wellcome Trust	
Loma willy (L.w.)	13	12		[Core Award	
				203141/Z/16/Z]	
Katharina Mara	40	0.7			
Katherine Wray	13	6,7		Haematology	
(K.W.)				Theme Oxford	
				Biomedical	
				Research	
				Centre	
Luzheng Xue (L.X.)	10,16	1,2,4,9,10,11,12	0000-0003-0042-1262	NIHR Oxford	
				BRC	
Hing Yuen Yeung	1	5	0000-0002-2353-7076	Wellcome	
(H.Y.Y.)				(204290/Z/16/Z)	
,				; Rosetrees	
				(R35579/AA002/	
				M85-F2)	
Zixi Yin (Z.Y.)	13	7		Chinese	
()				Academy of	
				Medical	
				Sciences	
				(CAMS)	
				Innovation Fund	
				for Medical	
				Science	
				(CIFMS), China	
				(grant number:	
				2018-I2M-2-002	
				and MRC Unit	
				Core	
Rebecca K Young	15	6,7			
(R.K.Y.)					
Jonathan Youngs	4,13	7,10			
(J.Y.)					
Ping Zhang (P.z.)	8	5	0000-0001-7063-7769		
Yasemin-Xiomara	1,7	2,3,5,6,8	0000-0002-3473-7129	BHF graduate	
Zurke (Y-X.Z.)				studentship FS/	
, ,				18/63/34184	

ISARIC4C Consortium	15	7	National Institute for Health Research (NIHR; award CO-CIN-01), the Medical Research Council (MRC; grant MC_PC_19059) , the NIHR Health Protection Research Unit in Emerging and Zoonotic Infections at University of Liverpool in partnership with Public Health England (PHE), in collaboration with Liverpool School of Tropical Medicine and the University of Oxford (NIHR award 200907)	The ISARIC WHO CCP-UK study protocol is available at https://isaric4c.net/protocols; study registry https://www.isrctn.com/IS RCTN66726260. This work uses data provided by patients and collected by the NHS as part of their care and support #DataSavesLives. We are grateful to the 2648 frontline NHS clinical and research staff and volunteer medical students who collected the data in challenging circumstances; and the generosity of the participants and their families for their individual contributions in these difficult times.
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 $^{{}^1} From\ CRediT\ taxonomy;\ https://www.cell.com/pb/assets/raw/shared/guidelines/CRediT-taxonomy.pdf$