



Supplementary Fig. S1. Highlighting decision-making matrix in deciding which secondary confirmation testing.

Supplementary Table S1. Breakdown of ASSD antibodies that were excluded due to missing data.

Samples excluded due to missing local data <i>Central testing results (n=84)</i>	
<i>ARS antibody</i>	<i>Number</i>
Jo1	1
PL7	3
PL12	4
EJ	3
OJ	0
KS	0
Zo	0
ARS Negative	73

Samples excluded due to missing central data <i>Local Testing Results (n=79)</i>	
<i>ARS Antibody</i>	<i>Number</i>
Jo1	29*
PL7	8
PL12	13*
EJ	2
OJ	4
KS	2
Zo	1
ARS Negative	21

Supplementary Table S2. Breakdown of the utilisation and performance of local testing based on the geographical location of assays randomly selected to undergo central immunoprecipitation.

Antibodies	Local (n)	Central (n)	Sensitivity (%)	Specificity (%)	PPV	NPV	Inter-rater agreement Weighted κ	
Europe n= 457								
					LIA = 76.4%	ELISA = 12.2%	IP = 2%	Other = 9.4%
ASSD (combined)	188	171	94.7	90.9	0.86	0.97	0.84 (95%CI 0.79-0.89)	
Jo1	120	124	96.7	100	1	0.99	0.98 (95%CI 0.96 – 1)	
Non-Jo1	68	47	80.9	92.7	0.56	0.98	0.61 (95%CI 0.5 – 0.72)	
PL-7	27	17	76.5	96.8	0.48	0.99	0.57 (95%CI 0.39-0.75)	
PL-12	19	24	84.2	98.2	0.67	0.99	0.73 (95%CI 0.58 – 0.88)	
EJ	9	13	66.7	98.4	0.46	0.99	0.53 (95%CI 0.28 – 0.79)	
OJ	3	0	n/a	99.3	n/a	1	-	
KS	2	2	0	99.6	0	0.99	-0.004 (95%CI -0.008 -0)	
Zo	0	0	-	-	-	-	-	
North America n = 48								
					LIA = 4.2%	ELISA = 45.8%	IP = 27.1%	Other = 22.9%
ASSD (combined)	36	35	100	92.3	0.97	1	0.95 (95%CI 0.84 - 1)	
Jo1	33	34	97.1	100	1	0.93	0.95 (95%CI 0.85 – 1)	
Non-Jo1	4	1	100	93.6	0.25	1	0.38 (95%CI -0.15 – 0.9)	
PL-7	0	0	-	-	-	-	-	
PL-12	3	1	100	95.7	0.33	1	0.48 (95%CI -0.11 - 1)	
EJ	1	0	n/a	97.9	0	1	-	
OJ	0	0	-	-	-	-	-	
KS	1	0	n/a	97.9	n/a	1	-	
Zo	0	0	-	-	-	-	-	
South America n= 30								
					LIA = 86.2%	ELISA = 10.3%	IP = 0%	Other = 3.5%
ASSD (combined)	26	24	100	66.7	0.92	1	0.76 (95%CI 0.45 - 1)	
Jo1	12	11	100	100	1	1	1 (95%CI 1 - 1)	
Non-Jo1	15	13	92.3	82.3	0.8	0.93	0.73 (95%CI 0.49 – 0.97)	
PL-7	3	2	100	94.6	0.67	1	0.78 (95%CI 0.37 - 1)	
PL-12	9	6	100	87.5	0.67	1	0.74 (95%CI 0.46 - 1)	
EJ	4	5	80	100	1	0.96	0.87 (95%CI 0.62 – 1)	
OJ	0	0	-	-	-	-	-	
KS	0	0	-	-	-	-	-	
Zo	0	0	-	-	-	-	-	
Japan n= 48								
					LIA = 2.1%	ELISA = 27.1%	IP = 70.8%	Other = 0%
ASSD (combined)	28	25	96	82.6	0.86	0.95	0.79 (95%CI 0.62-0.96)	
Jo1	9	10	90	100	1	0.97	0.93 (95%CI 0.81 - 1)	
Non-Jo1	19	15	93.3	84.8	0.74	0.97	0.73 (95%CI 0.53 – 0.93)	
PL-7	1	0	-	-	-	-	-	
PL-12	4	5	80	100	1	0.98	0.88 (95%CI 0.64-1)	
EJ	11	10	100	97.4	0.91	1	0.94 (95%CI 0.82- 1)	
OJ	1	0	n/a	97.9	n/a	1	-	
KS	2	0	n/a	95.8	n/a	1	-	
Zo	0	0	-	-	-	-	-	
India n= 41								
					LIA = 100%	ELISA = 0%	IP = 0%	Other = 0%
ASSD (combined)	16	15	93.3	92.3	0.88	0.96	0.84 (95%CI 0.68-1)	
Jo1	12	12	100	100	1	1	1 (95%CI 1-1)	
Non-Jo1	4	3	66.7	94.7	0.5	0.97	0.53 (95% CI 0.07 – 0.99)	
PL-7	1	2	0	97.4	0	0.95	-0.03 (95%CI -0.1 - 0.01)	
PL-12	3	2	50	94.9	0.33	0.97	0.36 (95%CI -0.2 - 0.92)	
EJ	0	0	-	-	-	-	-	
OJ	0	0	-	-	-	-	-	
KS	0	0	-	-	-	-	-	
Zo	0	0	-	-	-	-	-	

Supplementary Table S3. Double positive results for anti-ARS with the testing method used.

Antibodies double positive	Local/central test	Testing method
Anti-PL7/EJ	Local	Line blot
Anti-PL7/PL12	Local	Line blot
Anti-Jo1/KS	Local	Unknown
Anti-PL12/EJ	Local	Immunoprecipitation
Anti-PL7/PL12	Local	Line blot
Anti-Jo1/PL12	Local	Line blot
Anti-PL7/PL12	Central	ELISA

Supplementary Table S4. False positive and false negative rates of included ARS antibodies in the final analysis.

	Local Detection n, (%)	Central Detection n, (%)	False Positive Rate (%)	False Negative Rate (%)
Combined Testing n= 624				
ARS Combined	294 (47.1)	270 (43.3)	9.9	4.1
Jo1	186 (29.8)	191 (30.6)	0.2	3.1
Non-Jo1	110 (17.6)	79 (12.7)	7.9	15.2
PL7	32 (5.1)	21 (3.4)	2.8	28.6
PL12	44 (7.1)	33 (5.3)	2.7	15.2
EJ	29 (4.6)	24 (3.8)	1.5	16.7
OJ	4 (0.6)	0	0.6	-
KS	5 (0.8)	2 (0.3)	0.8	100
Zo	0	0	-	-
Local LIA n=419				
ARS Combined	184 (43.9)	166 (39.6)	10.3	4.8
Jo1	109 (26)	112 (26.7)	0.3	2.7
Non-Jo1	75 (17.9)	54 (12.9)	8.2	16.7
PL7	27 (6.4)	20 (4.8)	3.3	30
PL12	33 (7.9)	21 (5)	3.5	9.5
EJ	15 (3.6)	13 (3.1)	1.5	30.8
OJ	2 (0.5)	0	0.5	-
KS	1 (0.2)	1 (0.2)	0.2	100
Zo	0	0	-	-
Local ELISA n= 219				
ARS Combined	56 (60.9)	55 (59.8)	8.1	3.6
Jo1	47 (51.1)	47 (51.1)	0	0
Non-Jo1	9 (9.8)	8 (8.7)	3.6	25
PL7	1 (1.1)	0	1.1	-
PL12	4 (4.3)	5 (5.4)	1.1	40
EJ	3 (3.3)	2 (2.2)	1.1	0
OJ	1 (1.1)	0	1.1	-
KS	0	1 (1.1)	0	1
Zo	0	0	-	-
Local IP n = 56				
ARS Combined	31 (55.3)	28 (50)	14.3	3.6
Jo1	10 (17.9)	13 (23.2)	0	23.1
Non-Jo1	21 (37.5)	15 (26.8)	17.1	6.7
PL7	2 (3.6)	0	3.6	-
PL12	5 (8.9)	6 (10.7)	0	16.7
EJ	11 (17.9)	9 (16.1)	4.3	0
OJ	1 (1.8)	0	1.8	-
KS	3 (5.4)	0	5.4	-
Zo	0	0	-	-
Local Other Testing Methods n = 57				
ARS Combined	23 (40.4)	21 (36.8)	5.6	0
Jo1	19 (33.3)	19 (33.3)	0	0
Non-Jo1	5 (8.8)	2 (3.5)	5.5	0
PL7	2 (3.5)	1 (1.8)	1.8	0
PL12	2 (3.5)	1 (1.8)	1.8	0
EJ	0	0	-	-
OJ	0	0	-	-
KS	1 (1.8)	0	1.7	-
Zo	0	0	-	-

Supplementary Table S5. Contributing centres and site principal investigators for the CLASS Project.

Contributing centres of serum samples to CLASS Project	
Centres	Site PI
Ospedale Guglielmo da Saliceto, Piacenza, Italy	Elena Bravi
Keio University School of Medicine, Tokyo, Japan	Yuko Kaneko
Centro Hospitalar e Universitário de Coimbra, Portugal	Jose Antonio Pereira da Silva
Hospital General Universitario of Alicante, Spain	Vega Jovani
Hospital Universitario Ramon y Cajal, Madrid, Spain	Javier Bachiller-Corral
Hospital Universitario Marques de Valdecilla, IDIVAL, University of Cantabria Santander, Spain	Jose Cifrian
Hospital Clínico Universitario de Santiago de Compostela, Spain	Antonio Mera-Varela
University of Pittsburgh, USA	Siamak Moghadam-Kia
Instituto Nacional del Torax, Chile	Veronica Wolff
HP Metz, Hopital Belle-Ile, Metz, France	Julien Campagne
Centre de référence des Maladies auto-immunes rares, Hôpital de Hautepierre, Hôpitaux Universitaires de Strasbourg, France	Alain Meyer
ACURA Center for Rheumatic Diseases, Bad Kreuznach, Germany	Konstantinos Triantafyllias
University of Erlangen, Germany	Johannes Knitza
Department of Clinical Immunology and Rheumatology, SGPGIMS, Raebareli Road, Lucknow 226014, India	Latika Gupta Vikas Agarwal
Rabin Medical Center, Beilinson Hospital, and Sackler Faculty of Medicine, Tel Aviv University, Israel	Yair Molad
University of Bari, Italy	Florenzo Iannone Marco Fornaro
University and AO Spedali Civili, Brescia, Italy	Ilaria Cavazzana
University Clinic and AOU of Cagliari, Italy	Matteo Piga
IRCCS San Raffaele Hospital, Vita-Salute San Raffaele University, Italy	Giacomo De Luca
Participating centres within CLASS Project	
Centres	Site PI
Hospital María Ferrer, Argentina	Maria Laura Alberti
Fiona Stanley Hospital, Murdoch University, Notre Dame University, Perron Institute, Australia	Merrilee Needham
Clinique Universitaire Saint Luc, Belgium	Frederic Houssiau
Faculdade de Medicina, Universidade de Sao Paulo, Brazil	Samuel Shinjo
University of Western Ontario, Canada	Pari Basharat
Peking Union Medical College Hospital, Peking Union Medical College & Chinese Academy of Medical Sciences, China	Qian Wang
Universidad Nacional de Colombia/Fundacion Santa fe de Bogota, Colombia	Gerardo Quintana
CHU Lyon, France	Laure Gallay
Bichat-Claude Bernard Hospital, France	Esther Ebstein
Medical Center Baden-Baden, Germany	Christopher Fiehn
Helios Gesundheit, Germany	Eugene Feist
University Medical Center Dusseldorf, Germany	Matthias Schneider
Johann Wolfgang Goethe-Universität, Frankfurt, Germany	Ulrich Drott
University Medical Center of Freiburg, Germany	Reinhard Edmund Voll
University of Debrecen, Hungary	Katalin Danko
St John's National Academy of Medical Sciences, Bangalore, India	Vineeta Shobha
Nizam's Institute of Medical Sciences, India	Liza Rajasekhar
Clinical Medicine, University of Ancona, Italy	Maria Giovanna Danieli
ASST Papa Giovanni XXIII, Bergamo, Italy	Massimiliano Limonta
University of Florence, Italy	Marco Matucci-Cerinic
University of Florence, Italy	Giacomo Emmi
University of Florence, Dept. Experimental and Clinical Medicine, Florence, Italy	Paola Parronchi
U.O.C. Malattie Respiratorie Dip. Scienze Mediche, Chirurgiche e Neuroscienze, Università di Siena Policlinico "Le Scotte", Italy	Elena Bargagli

Participating centres within CLASS Project

Centres	Site PI
Fondazione IRCCS Istituto Neurologico Carlo Besta, Italy	Lorenzo Maggi
Azienda Ospedaliera Universitaria di Modena, Italy	Marco Sebastiani
University of Milano Bicocca, San Gerardo Hospital, Monza, Italy	Alberto Pesci
Università del Piemonte Orientale UPO, Novara, Italy	Pier Paolo Sainaghi
University of Perugia, Italy	Elena Bartoloni-Bocci
Azienda Ospedaliera Universitaria Pisana, Pisa, Italy	Simone Barsotti
Graduate School of Medicine, Kyoto University, Kyoto, Japan	Ran Nakashima
IRCCS ASMN Reggio Emilia, Italy	Nicolò Pipitone
University La Sapienza and Policlinico Umberto I, Rome, Italy	Valeria Ricciari
Ospedale San Camillo, Roma, Italy	Giandomenco Sebastiani
Ospedale “ Scarlato” Scafati, Italy	Salvatore Scarpato
Azienda Ospedaliera Universitaria Senese - Ospedale Santa Maria alle Scotte, Siena, Italy	Luca Cantarini
Ospedale Santa Chiara, Trento, Italy	Alvise Berti
ASST Bergamo Ovest - Treviglio Hospital, Italy	Paolo Luigi Colombelli
University Hospital of Cattinara, Trieste, Italy	Paola Tomietto
Ospedale Mauriziano, Turin, Italy	Claudia Lomater
Santa Maria della Misericordia Hospital and University of Udine, Italy	Luca Quartuccio
Verona University, Italy	Giovanni Orsolini
Tosei General Hospital, Seto, Japan	Yasuhiro Kondo
Tokyo Medical and Dental University, TMDU, Bunkyo-ku, Japan	Naoki Kimura
OPD Hospital Civil Dr. Juan I. Menchaca, Mexico	Monica Vazquez-Del Mercado
Oslo University Hospital, Norway	Helena Andersson
Centro Hospitalar Lisboa Norte, Portugal	Joao Eurico Cabral de Fonseca
Hospital Clínic of Barcelona/IDIBAPS - University of Barcelona, Spain	Sergio Prieto-González
Hospital Universitario Dr. Negrín, Las Palmas, Spain	Martín Gerardo Greco Merino
Hospital Universitario de la Princesa, IIS-Princesa, Madrid, Spain	Santos Castaneda
Hospital General Universitario Gregorio Marañón, Madrid, Spain	Julia Martinez Barrio
Hospital Universitario La Paz, Madrid, Spain	Laura Nuno
Hospital Universitari Mutua Terrassa, Terrassa, Spain	Ignasi Rodriguez Pintó
Hospital Universitario Araba, Spain	Jaime Calvo Alén
Hospital Universitario Miguel Servet, Zaragoza, Spain	Luis Sáez-Comet
Lund University and Skåne University Hospital Malmö, Sweden	Giovanni Cagnotto
Akdeniz University, School of Medicine, Turkey	Erkan Alpsoy
University of Manchester, UK	Hector Chinoy
Michigan Medicine, University of Michigan, USA	Elena Schiopu
Emory University, USA	Prateek Gandiga
University of Kansas Medical Center, USA	Mazen Dimachkie
University of California, Los Angeles, USA	Christina Charles-Schoeman
Vanderbilt University Medical Center, USA	Erin Wilfong
Northwell Health, USA	Galina Marder
North Bristol NHS Trust, UK	Harsha Gunawardena
Oregon Health and Science University, USA	Daniela Ghetie