Supplementary Table. Demographic, clinical, radiological, laboratory and neuropathological characteristics of forty-three patients with definite, probable and possible COVID-19 infection with central and peripheral neurological syndromes.

Table S1a: Ten patients with encephalopathy (delirium and psychosis)

Table S1b: Twelve patients with inflammatory CNS syndromes (encephalitis, para- and post- infectious) (2 parts)

Table S1c. Demographic, clinical, radiological, laboratory and characteristics of eight individuals with possible and

definite COVID-19 infection with ischaemic stroke

Table S1d. Demographic, clinical, radiological, laboratory and characteristics of individuals with possible and definite

COVID-19 infection with peripheral neurological syndromes

Table S1e. Five miscellaneous and uncharacterised patients.

Abbreviations used in Tables. CADASIL - Cerebral autosomal dominant arteriopathy with subcortical infarcts and leukoencephalopathy; TIA – transient ischaemic attack; ICU – intensive care unit; N/A – not applicable; Hb – haemoglobin; MCV - mean corpuscular volume; Lymphs - lymphocytes; Neuts - neutrophils; Plt - platelets; ESR - erythrocyte sedimentation rate; CRP - C-reactive protein; NR - no result; TSH - thyroid stimulating hormone; NMDA receptor - N-methyl-D-aspartate receptor; CSF - cerebrospinal fluid; PCR - polymerase chain reaction; CT - computed tomography; MRI – magnetic resonance imaging; CXR – chest X-ray; CTPA – CT pulmonary angiogram; ARDS – acute respiratory distress syndrome; EEG - electroencephalogram; IV - intravenous; IVMP - intravenous methylprednisolone; CPAP – continuous positive airway pressure; ADEM – acute demyelinating encephalomyelitis; MGUS – monoclonal gammopathy of undetermined significance; ANA - anti-nuclear antibodies; ANCA - anti-neutrophil cytoplasmic antibodies; ENA – extractable nuclear antigens; IgG – Immunoglobulin G; GAD – glutamic acid decarboxylase; DPPX - dipeptidyl-peptidase-like protein 6; LGi1 - leucine-rich glioma-inactivated 1; CASPR2 - contactinassociated protein-like 2; MOG – myelin oligodendrocyte glycoprotein; AOP4 – aquaporin-4; FDG PET – fluorodeoxyglucose positron emission tomography; IVIG - intravenous immunoglobulin; JPS - joint position sense; HTLV 1 and 2 - human T-cell leukaemia virus types 1 and 2; EMG - electromyogram; ICP - intracranial pressure; DVT deep vein thrombosis; PPM - permanent pacemaker; AF - atrial fibrillation, MCA - middle cerebral artery; ACA anterior cerebral artery; PCA - posterior cerebral artery; mRS - modified Rankin Scale; NHISS - National Institute for Health Stroke Scale; CTA - CT angiogram; PE pulmonary embolism; APTT - activated partial thromboplastin time: IgM – immunoglobulin M: LDL – low density lipoprotein: HDL – high density lipoprotein: GBS – Guillian Barre syndrome; Hep E – hepatitis E; NAD – no abnormality detected; GORD – gastro oesophageal reflux disease; AML – acute myeloid leukaemia; QDS – four times daily; EVD – external ventricular drain

Standard viral PCR – herpes simplex viruses, varicella zoster, enterovirus +/- adenovirus, cytomegalovirus, Epstein-Barr Virus, parechovirus depending upon NHS trust/centre; N-Gene PCR (unvalidated test) for SARS-CoV-2 testing in CSF only where specified.

Supplementary Table. Demographic, clinical, radiological, laboratory and neuropathological characteristics of forty-three patients with definite, probable and possible COVID-19 infection with central and peripheral neurological syndromes.

S1a: Ten patients with encephalopathy (delirium and psychosis)

Patient	1	2	3	4	5	6	7	8	9	10
Age (years)	65	72	59	58	52	39	55	68	50	57
Age (years)	05	12	57	50	52	57	55	00	50	57
Gender (M/F)	F	М	F	М	F	F	F	М	F	М
Ethnicity	White	White	Black	Black	White	Asian	White	Black	Black	Black
Final neurologi- cal diagnosis	Hypoactive de- lirium	Hypoactive delirium	Delirium	Delirium	Delirium	Delirium	Delirium and psychosis	Hyperactive delirium	Generalised tonic clonic seizures with encephalopa- thy	Encephalopathy with myelopathy and ataxia
Past medical history	CADASIL; pre- vious right oc- cipital stroke; TIA	Bladder can- cer; nephrec- tomy; hyper- cholesterolae- mia	Hypertension; diabetes	Prostate can- cer in remis- sion; muscu- loskeletal back pain	None	Hypertension	None	None	None	Hypertension; di- abetes; asthma
Initial COVID- 19 symptoms	Cough; fever	Confusion; fe- ver	Diarrhoea; vomiting; fe- ver; cough; confusion	Confusion; cough; fever; dysgeusia	Fever; lethargy; confusion	Fever; cough; dyspnoea	Cough; dysp- noea; fever; myalgia; anos- mia; hypoge- usia	Fever	Cough; household contacts with COVID-19 symp- toms	Cough; dyspnoea; fever; myalgia
Days from onset of	of COVID-19 symp	otoms to:								
1) Hospital ad- mission	15	4	7	4	3	7	1st 14; 2nd 17	1	21	6
2) Onset of neu- rological symp- toms	14	1	3	1	-1	19	17	-4	21	6
3) ICU admis- sion	N/A	6	N/A	N/A	N/A	7	N/A	N/A	N/A	N/A
Main reason for hospital admis-	Neurological	Both	Neurological	Neurological	Neurological	Respiratory	1st Respira- tory; 2nd Neu- rological	Neurological	Neurological	Both

sion (neurologi- cal vs respira- tory symptoms)										
Security of COVID-19 di- agnosis (defi- nite, probable, possible)	Definite	Definite	Definite	Definite	Probable	Definite	Definite	Definite	Probable	Definite
Severity of COVID-19 in- fection	Mild	Critical	Mild	Mild	Mild	Critical	Severe	Mild	Mild	Severe
Duration of ICU stay (days)	N/A	13	N/A	N/A	N/A	16	N/A	N/A	N/A	N/A
Initial neuro- logical symp- toms	Fluctuating confusion; re- versal of sleep- wake cycle	Confusion; malaise; loss of appetite	Fluctuating confusion	Confusion; nonsensical speech; repeti- tive behav- iour; disorien- tation; delu- sional thoughts; headache	Fluctuating con- sciousness; de- lirium	Delirium; hallu- cinations about experiences in countries not previously vis- ited; reversed sleep/wake cy- cle	Confusion; agitation; per- secutory delu- sions; visual hallucinations; combative be- haviour; head- aches	Cognitive im- pairment; gait disturbance; two falls	Seizures	Double inconti- nence; progres- sive lower > up- per limb weak- ness; disorienta- tion; word find- ing difficulties
Key neurologi- cal signs	Disorientated to time and place; impaired in- sight, brady- phrenia; polyminimyo- clonus; old left homonymous hemianopia	Cognitive im- pairment; in- creased limb tone; brisk re- flexes	Fluctuating at- tention and cognition; bradyphrenia; dyspraxia.	Bilateral in- tention tremor; heel- shin ataxia	Cognitive im- pairment; re- duced verbal fluency	Cognitive im- pairment	No focal signs	Disorientation; intermittent agi- tation; unable to follow com- mands; speak- ing a few words only; bilateral extensor plan- tars	Post-ictal drowsi- ness and transient disorientation only	Pyramidal tract signs; extensor plantars; mild up- per limb dysme- tria
Initial laboratory	results at or neare	est to time of neu	rological sympto	m onset						
Hb (g/L; 130- 170)	130	147	118	162	107	86	113	137	113	172
MCV (fL; 80- 99)	84.4	95.4	70.3	79.4	89	89.8	85	73.9	97.2	93.3
Lymphs (x10^9/L; 1.2- 3.65)	1.47	0.43	2.31	0.26	0.91	1.16	1.42	1.39	4.48	0.69

Neuts (x10^9/L; 2-7.5)	5.97	4.09	5.09	9.93	5.74	9.52	4.26	3.29	4.65	2.29
Plt (x10^9/L; 150-400)	479	106	276	191	158	599	454	123	379	81
ESR (mm/hr; 1- 20)	NR	NR	NR	NR	61	143	NR	NR	NR	NR
CRP (mg/L; 0- 5.0)	58.3	64.6	38.1	168.1	56	14.8	37.9	38	8.9	85.5
Fibrinogen (g/L; 1.5-4.0)	NR	NR	NR	NR	4.6	8	7.82	NR	NR	NR
D-Dimer (µg/L; 0-550)	1190	1730	NR	970	NR	2430	1200	NR	NR	1410
Ferritin (µg/L; 30-400)	721	1306	425	3285	393	969	1291	NR	281	1563
Phospholipid an- tibodies	NR	NR	NR	NR	Negative	NR	NR	NR	NR	NR
Lupus anticoag- ulant	NR	NR	NR	NR	NR	NR	NR	NR	NR	NR
Other relevant results				Free T4 24.2 (normal range 12-22), TSH 0.07 (0.27- 4.20)			Significant negative re- sults: neuronal and NMDA receptor anti- bodies			
CSF:			1							
White cell count (cells/µL; 0-5)	NR	NR	3	2	NR	NR	<1	3	<1	<1
Protein (g/L; 0.13-0.45)	NR	NR	0.52	Insufficient	NR	NR	0.18	0.4	0.43	0.36

Glucose (mmol/L; 2.2- 4.2) (blood glu-	NR	NR	NR	3.17 (5.9)	NR	NR	3.5 (5.0)	3.7 (5.3)	NR	3.7 (6.0)
cose) (mmol/L; 3.9-5.8)										
Other			CSF culture and viral PCR negative	CSF culture and viral PCR negative			CSF culture and viral PCR negative	CSF culture and viral PCR nega- tive	Oligoclonal bands negative; CSF culture and viral PCR negative	Matched oligo- clonal bands in serum and CSF; CSF culture and viral PCR nega- tive
Micro/virology	Blood, urine and sputum cul- tures negative	Blood and urine cultures negative	Blood cultures negative	Blood and urine cultures negative			Blood and urine cultures negative		Screening for other respiratory viruses negative	Blood, urine, stool cultures all negative
Imaging	l	I	I	l	l	l	1	I	1	
Neuroaxis	CT head - no acute changes	CT head - small vessel disease, no acute changes	CT head - small vessel disease	MRI brain normal	CT head - cal- cification con- sistent with Fahr's disease; MRI brain – consistent with CT small foci of restricted diffu- sion in the both centrum semio- vale	MRI brain - fo- cal non-specific T2/FLAIR-hy- perintensities within globi pallidi bilater- ally	MRI brain normal	MRI brain normal	MRI brain normal	MRI brain - mild to moderate small vessel disease, MRI spine - mul- tilevel degenera- tive disc disease only
Chest	CXR - bilateral infiltrates sug- gestive of COVID-19	CXR - pro- gressive bilat- eral, symmet- rical nodular infiltrates con- sistent with COVID-19	CXR – ill-de- fined bibasal infiltrates and peripheral air- space opacifi- cation left lower zone; indeterminate for COVID- 19	CXR - normal	CXR – periph- eral airspace opacification consistent with COVID-19	CTPA – exten- sive bilateral ground glass ap- pearance and consolidation consolitent with COVID-19 ARDS	CT chest – bi- lateral sub- pleural ill-de- fined ground- glass opacities and interlobu- lar septal thickening consistent with COVID- 19	CXR - bibasal infiltrates; CT chest - ground glass changes	CXR - ill-defined peripheral air- space opacifica- tion in both lungs, in keeping with probable COVID- 19	CXR - Bilateral lower lobe opaci- ties consistent with COVID-19
Other relevant in	vestigations	I	I	1	I	I		I	1	
Neurophysiol- ogy							EEG within normal limits	EEG within normal limits		

Treatment										
For neurological diagnosis	Supportive	Supportive	Supportive	Supportive	Supportive	Melatonin	Haloperidol; risperidone	1g IVMP 3 days for initial suspi- cion of autoim- mune encephali- tis	IV lorazepam acutely for sei- zures; levetirace- tam	Supportive
For COVID-19 infection	None required	CPAP	None required	None required	None required	Intubation and ventilation	Oxygen	None required	None required	Oxygen
Outcomes										
Total duration of hospital admis- sion (days)	8	39	14	4	12	20	23	Ongoing	2	12
Outcome at last follow up	Complete re- covery; dis- charged home	Complete re- covery of con- fusion; dis- charged to re- habilitation centre	Complete re- covery; dis- charged home	Complete re- covery; dis- charged home	Complete recov- ery; discharged home	Incomplete re- covery; ongoing cognitive im- pairment; dis- charged home	Significant re- covery; im- proving; dis- charged home	Ongoing admis- sion; receiving treatment	Complete recov- ery; discharged home	Complete recov- ery; discharged home

D (11	10	10	14	15	16
Patient	11	12	13	14	15	16
Age (years)	65	66	52	60	59	52
Gender (M/F)	F	F	М	М	F	М
Ethnicity	Black	White	Asian	Asian	Asian	White
Final neurological diagnosis	Possible post-infectious Encephalitis (presumed autoimmune)	Encephalitis	ADEM (with haemorrhage)	ADEM (with haemorrhage)	ADEM (with necrosis and haemorrhage)	ADEM (with haemorrhage) and acute demyelinating poly- radiculoneuropathy
Past medical his- tory	Possible Alzheimer's dis- ease; osteoarthritis	Hypertension; hypothy- roidism; hysterectomy; osteoarthritis; degenera- tive spine disease	Asthma	Diabetes; hypertension	Aplastic anaemia; MGUS; treated breast cancer; hyper- tension; non-alcoholic fatty liver disease; hypercholes- terolaemia	None
Initial COVID-19 symptoms	Cough; fever; myalgia; diarrhoea	Fever	Cough; myalgia; dyspnoea; hypoxia	Fatigue; myalgia; fever; dyspnoea; hypoxia	Cough; chills; lethargy; my- algia	Fever; hypoxia
Days from onset of Co	OVID-19 symptoms to:		I			
1) Hospital admis- sion	11	1	10	8	10	-4
2) Onset of neuro- logical symptoms	6	1	22	27	10	-6
3) ICU admission	N/A	N/A	10	8	10 (for deterioration in con- sciousness)	1
Main reason for hospital admission (neurological vs respiratory symp- toms)	Neurological	Neurological	Respiratory	Respiratory	Neurological	Neurological
Security of COVID- 19 diagnosis	Definite	Definite	Definite	Definite	Definite	Definite

Severity of COVID- 19 infection	Severe	Mild	Critical	Critical	Mild	Critical
Duration of ICU stay (days)	N/A	N/A	35	35	9	14
Initial neurological symptoms	'Shaking'; confusion; hal- lucinations; reported be- ing 'cross-eyed'; dysar- thria	Confusion; single general- ised seizure	Slow to wake in ICU	Slow to wake in ICU	Recurrent fleeting episodes of vacant staring and speech arrest; generalised tonic- clonic seizures; headache; low conscious level	Headache; back pain; vomit- ing; progressive limb weak- ness
Key neurological signs	Perseveration; opsoclo- nus; convergence spasm; bilateral ocular-facial synkinesis; dysarthria; orofacial apraxia; general- ised stimulus-sensitive myoclonus	Low conscious level post- ictally lasting 48 hours; brisk reflexes; extensor plantars	Low conscious level; with- drawal to pain; hyperre- flexia and clonus	Low conscious level; open- ing eyes to voice; with- drawal to pain; right exten- sor plantar	Left pupil unreactive at na- dir; left extensor plantar	Flaccid four limb weakness, proximal > distal; facial and neck weakness; areflexia; ex- tensor plantars, normal sensa- tion; ophthalmoplegia day 3
Initial laboratory resu	ilts at or nearest to time of 1	neurological symptom onset				
Hb (g/L; 130-170)	113	119	86	93	105	155
MCV (fL; 80-99)	95.5	81.5	73.1	96	90	91
Lymphs (x10^9/L; 1.2-3.65)	1.68	0.4	3.07	0.95	0.3	1.07
Neuts (x10^9/L; 2- 7.5)	11.02	5.58	13.96	12.79	2.7	5.01
Plt (x10^9/L; 150- 400)	759	193	760	370	29 (premorbid baseline 33)	254
ESR (mm/hr; 1-20)	76	NR	NR	NR	NR	22
CRP (mg/L; 0-5.0)	9.1	14	91.5	149	8.1	1
Fibrinogen (g/L; 1.5- 4.0)	5.15	3.29	6.5	NR	4.08	NR
D-Dimer (normal range for assay)	1800 µg/L (0-550)	1599 ng/mL (0-230) (day 15, post-IVIG)	80,000 μg/L (0-550)	3330 mg/L (250-750)	2033 mg/L (250-750)	NR
Ferritin (µg/L; 30- 400)	261	183	2661	533	98	287

Phospholipid anti- bodies	NR	NR	NR	NR	NR	NR
Lupus anticoagulant	NR	NR	NR	NR	NR	NR
Other relevant re- sults	ANA positive 1:1280, thyroid peroxidase anti- bodies positive; paraproteins: 3g IgG lambda, 4g IgG kappa; Significant negatives: neuronal and ganglioside antibodies; antibodies to glycine receptor, GAD, DPPX, NMDA-receptor, LGI1 and CASPR2; ANCA; ENA; comple- ment	Significant negatives: ANA; ANCA; neuronal antibodies; antibodies to LGI1, CASPR2, NMDA receptor GAD, thyroid pe- roxidase.	ANA positive 1:1280; ele- vated IgG; Significant nega- tives: neuronal antibodies, antibodies to NMDA-recep- tor, LGI1, CASPR2, MOG, AQP4, GAD	ANA and ANCA negative		ANA and ANCA negative
CSF						
White cell count (cells/µL; 0-5)	1	3	1	1	4	0
Protein (g/L; 0.13- 0.45)	0.22	1.0	0.38	0.22	2.34	1.01
Glucose (mmol/L; 2.2-4.2) (blood glu- cose) (mmol/L; 3.9- 5.8)	3.2 (4.9)	3.5	4.01 (5.37)	5.5	NR	3.6 (6.4)
Other	Matched oligoclonal bands in serum and CSF; neuronal antibodies nega- tive; antibodies to NMDA receptor and GAD nega- tive; CSF culture and viral PCR negative	Oligoclonal bands nega- tive; CSF culture and viral PCR negative, including SARS-CoV-2	Oligoclonal bands negative; Antibodies to NMDA re- ceptor negative; CSF cul- ture and viral PCR negative	CSF culture and viral PCR negative including SARS- CoV-2	Opening pressure 28 cm H20; CSF viral PCR nega- tive including SARS-CoV-2	CSF viral PCR ; negative in- cluding SARS-CoV-2
Micro/virology	Blood cultures negative	Blood cultures negative		Blood and urine cultures negative; screening for other respiratory viruses negative		
Imaging	1	1		negative	1	
Neuroaxis	MRI brain normal	MRI brain - T2 hyperin- tense signal changes in upper pons, limbic lobes,	MRI brain - Multiple clus- ters of lesions in the deep cerebral white matter. Cyst- like areas of varied sizes,	MRI brain - Multifocal and confluent areas of signal change in the cerebral hemi- spheric white matter with	Initial CT head - possible early brainstem swelling, re- peat CT day 2 showed hypo- density of brainstem and	MRI brain - Multifocal con- fluent lesions in internal and external capsules, splenium

		medial thalami and sub- cortical cerebral white matter. The limbic lobe involvement was symmet- rical and included punc- tate diffusion abnormali- ties. No micro-haemor- rhages or enhancement.	some with haemorrhagic foci and peripheral rims of restricted diffusion were shown within these clusters.	extensive microhaemor- rhages in the subcortical re- gions.	thalami; MRI brain (day 6) Extensive, confluent and largely symmetrical areas throughout brain stem, lim- bic and insular lobes, super- ficial subcortical white mat- ter and deep grey matter. Clusters of microhaemor- rhages, restricted diffusion and peripheral rim enhance- ment.	and deep white matter of cere- bral hemispheres. Over 5 days, these lesions increased in size and showed multiple microhaemorrhages and ex- tensive prominent medullary veins. Components of brachial and lumbosacral plexus showed increased signal and enhancement without overt swelling.
Chest	CT chest - severe COVID-19 related changes	CT chest suggestive of COVID-19.	CXR - bilateral, lower zone predominant, ill-defined air- space opacification, con- sistent with COVID-19	CXR - bilateral subpleural airspace infiltrates sugges- tive of COVID-19	CXR - bibasal atelectasis but no consolidation. Mild pul- monary venous congestion + mild bilateral pleural effu- sions	CT chest - bilateral infiltrates
Other imaging	FDG PET CT – no malig- nancy					
Other investigations	1					
Neurophysiology	EEG normal, no correlate for movements		EEG - non-specific cortical dysfunction, mild diffuse encephalopathy			NCS - moderately severe acute demyelinating poly- radiculoneuropathy; EEG - encephalopathy, no epilepti- form discharges
Treatment						
For neurological di- agnosis	1g IVMP 3 days, then oral prednisolone taper; le- vetiracetam; clonazepam	1g IVMP 3 days then oral prednisolone taper, IVIG	Supportive	1g IVMP 3 days	Intubation and ventilation; levetiracetam, aciclovir and ceftriaxone, dexamethasone	Intubation and ventilation; 1g IVMP 5 days; IVIG
For COVID-19 in- fection	Oxygen	None required	Intubation and ventilation; renal replacement	Intubation and ventilation	Intubation for low conscious level rather than respiratory reasons	Oxygen; intubation and venti- lation for type 2 respiratory failure
Other					Platelet infusions for throm- bocytopenia	
Outcomes						I
Total duration of hospital admission (days)	22	28	Ongoing	Ongoing	10	Ongoing
Outcome at last fol- low up	Incomplete recovery; slow improvement but on- going myoclonus; dis- charged home	Incomplete recovery; on- going cognitive impair- ment; discharged home	Incomplete recovery; im- proving	Incomplete recovery; im- proving	No response to treatment; died	Incomplete recovery; improv- ing; ongoing rehabilitation

Table S1b: Twelve patients with inflammatory CNS syndromes (encephalitis, para- and post- infectious) (part 2 of 2)

Patient	17	18	19	20	21	22
Age (years)	47	54	60	33	27	48
Gender (M/F)	F	F	F	F	F	М
	1	Ĩ	1	Ĩ	Ĩ	111
Ethnicity	Other	Mixed	Black	White	Asian	White
Final neurological diagnosis	ADEM (with haemorrhage)	ADEM	ADEM	ADEM with myelitis	ADEM with myelitis	Post-infectious myelitis
Past medical his- tory	asthma	Hypertension; polycystic ovarian syndrome	Hypertension; diabetes	None	None	Hypertension; diabetes
Initial COVID-19 symptoms	Cough; fever; dyspnoea	Cough; fever; dysgeusia; truncal rash	Cough; dyspnoea; diar- rhoea; fever	Fever	Cough; fever; anosmia; dysgeusia	Cough; dyspnoea; fever
Days from onset of C	COVID-19 symptoms to:					1
-						
1) Hospital admis- sion	8	23	4	4	21	1st: day 3; 2nd: day 7; 3rd: day 21
1) Hospital admis-	8	23	4	4	21 8	
1) Hospital admis- sion 2) Onset of neuro-						day 21
 Hospital admission Onset of neurological symptoms ICU admission Main reason for hospital admission (neurological vs respiratory symp- 	8 11 (for deterioration in con-	14	18	2 4 (for deterioration in con-	8	day 21 19
 Hospital admission Onset of neurological symptoms ICU admission ICU admission Main reason for hospital admission (neurological vs respiratory symptoms) Security of COVID-19 diagno- 	8 11 (for deterioration in con- sciousness)	14 N/A	18	2 4 (for deterioration in con- sciousness)	8 N/A	day 21 19 N/A 1st Respiratory ; 2nd Respira-
 Hospital admission Onset of neurological symptoms ICU admission ICU admission Main reason for hospital admission (neurological vs respiratory symptoms) Security of 	8 11 (for deterioration in con- sciousness) Neurological	14 N/A Neurological	18 4 Respiratory	2 4 (for deterioration in con- sciousness) Neurological	8 N/A Neurological	day 21 19 N/A 1st Respiratory ; 2nd Respira- tory; 3rd Neurological

Initial neurological symptoms	Subacute left sided numb- ness and weakness; head- ache; vomiting; reduced conscious level	Unsteadiness; left sided limb weakness; slurred speech; fatigue; falls	Slow to wake in ICU	Headache; confusion; re- duced conscious level	Sensory symptoms in feet and right hand; difficulty with balance and walking	Numbness of hands and feet; band of itching sensation at level of the umbilicus; un- steady gait
Key neurological signs	Dense left hemiparesis; re- duced sensation on left	Drowsy; slow to respond; dysarthric; trunk and limb ataxia; broad base stand- ing; unable to walk; left- sided pyramidal weak- ness; bilateral extensor plantars	Extensor posturing of limbs	Low conscious level; brain- stem breathing pattern; at worst locked-in; areflexic	Spastic gait, broad based; increased tone with ankle clonus; pyramidal weak- ness; loss of pinprick and fine touch in feet; no sen- sory level	Weakness of hip flexion; brisk reflexes; extensor plantars; vi- bration impaired to costal margins, JPS to ankles, pin- prick to T10; sensory ataxia
Initial laboratory res	ults at or nearest to time of n					
Hb (g/L; 130-170)	130	139	68	112	130	157
MCV (fL; 80-99)	84.2	75.1	93.1	93.7	82.4	89.2
Lymphs (x10^9/L; 1.2-3.65)	0.92	2.3	1.74	1.79	1.81	2
Neuts (x10^9/L; 2- 7.5)	7.26	6.2	10.12	8.52	2.35	3.37
Plt (x10^9/L; 150- 400)	271	345	336	174	249	278
ESR (mm/hr; 1-20)	58	NR	88	43	12	NR
CRP (mg/L; 0-5.0)	34	18.5	169.8	7.6	1.2	<5
Fibrinogen (g/L; 1.5-4.0)	NR	5.7	NR	6.92	NR	3.25
D-Dimer (normal range for assay)	1160 µg/L (0-550)	NR	>8000 ng/mL (<500)	2210 μg/L (0-550)	NR	NR
Ferritin (µg/L; 30- 400)	92	NR	521	998	NR	473
Phospholipid anti- bodies	NR	Negative	NR	Negative	NR	NR
Lupus anticoagulant	NR	Positive	Negative	Positive	NR	NR

Other relevant re- sults	Significant negatives: neu- ronal antibodies; antibodies to AQP4 and MOG		ANA and ANCA nega- tive	Significant negatives: ANA, ENA, ANCA; anti- bodies to AQ4, MOG	ANA and ANCA negative	
CSF:						
White cell count (cells/µL; 0-5)	Not done (mass effect on imaging).	19 (10% polymorphs, 90% lymphocytes),	<5	54 (95% polymorphs)	4	10 (lymphocytes),
Protein (g/L; 0.13- 0.45)		0.33	0.36	0.31	0.52	0.7
Glucose (mmol/L; 2.2-4.2) (blood glu- cose) (mmol/L; 3.9- 5.8)		4.1 (5.5)	8.4 (12.5)	7.2 (13.5)	3.1 (4.6)	5.6
Other		Oligoclonal bands nega- tive; CSF culture - scanty growth of Staphylococcus capitis - likely contami- nant	Oligoclonal bands nega- tive; CSF viral PCR in- cluding SARS-CoV-2 negative	Oligoclonal bands negative; antibodies to MOG, AQP4, NMDA receptor, CASPR2 and LGi1 all negative; CSF culture and viral PCR nega- tive including SARS-CoV-2	CSF culture negative; anti- bodies to AQP4 and MOG pending at time of publica- tion	Matched oligoclonal bands in serum and CSF; CSF culture and viral PCR negative
Micro/virology	Blood and urine cultures negative; screen for other respiratory viruses negative; Brain biopsy - viral PCR negative including SARS- CoV-2; mycoplasma PCR negative.		Blood, urine, sputum cultures negative; HTLV1 negative	Tic-borne encephalitis vi- rus, Lyme, returning travel- ler screen (Porton Down) all negative	HTLV1 serology negative	Blood and urine cultures neg- ative, HTLV 1/2 negative, syphilis negative
Imaging			I	I		
MRI neuroaxis	Severe right hemispheric vasogenic oedema with a leading edge on contrast im- aging. Smaller areas of T2 hyperintense changes in the left hemisphere. Marked mass-effect with 10mm left- wards midline shift, and mild subfalcine herniation	Multiple large lesions with peripheral rim re- striction in periventricular white matter of both cere- bral hemispheres	Multifocal lesions with diffusion changes in periventricular white matter and corpus callo- sum.	Initial MR showed multifo- cal lesions in lower brain- stem, medial temporal lobes and cerebral white matter, some of which showed re- stricted diffusion. 3 days later, the brainstem lesions coalesced and extensive in- tramedullary lesions with swelling involving grey and white matter of the spinal cord appeared. At 3 week follow-up, these lesions persisted but without swell- ing or oedema.	Diffuse ill-defined conflu- ent T2 hyperintensity in- volving the white matter of the cerebral hemispheres, largely along the corticospi- nal tracts. Small focal area of diffusion change in the left motor cortex. Ill-de- fined intramedullary lesion without swelling in the co- nus medullaris.	MRI brain normal; MRI tho- racic spine - a patchy area of intramedullary high signal in the dorsal cord primarily at T5-6 and T10-11 and down to the conus with no enhance- ment with contrast. Felt to be consistent with post-infectious myelitis.
Chest	CTPA – changes at lung ba- ses that could be compatible with COVID-19 infection	CXR - normal	CT chest - bilateral pul- monary infiltrates	CT chest - not typical for COVID-19	NR	CXR - patchy infiltrates; Fur- ther CXR - pneumonia

Other relevant inves	tigations					
Neurophysiology			EEG – diffuse encepha- lopathy	EEG - diffuse encephalopa- thy		Normal nerve conduction studies and EMG
Histology	Brain biopsy - histology consistent with ADEM					
Treatment given	I					
For neurological di- agnosis	Intubation for low con- scious level; right hemicra- niectomy; 1g IVMP 5 days, then oral prednisolone; IVIG.	1g IVMP 3 days, then oral prednisolone	1g IVMP 3 days, then oral prednisolone taper	Intubation for low con- scious level; ICP bolt; lum- bar drain; 1g IVMP 3 days then oral prednisolone	None	1g IVMP 3 days
For COVID-19 in- fection	Oxygen	None required	Intubation and ventila- tion; renal replacement	None required	None required	None required
Other						Antibiotics for secondary bac- terial pneumonia
Outcomes						
Total duration of hospital admission (days)	Ongoing	Ongoing	Ongoing	Ongoing	0 (Outpatient management)	1st and 2nd: 1 day each; 3rd: 9 days
Outcome at last fol- low up	Incomplete recovery; im- proving	Incomplete recovery; im- proving	Incomplete recovery; improving	Incomplete recovery; im- proving	Complete recovery; at home	Incomplete recovery; improv- ing; ongoing rehabilitation

Table S1c. Demographic, clinical, radiological, laboratory and characteristics of eight individuals with possible and definite COVID-19 infection with ischaemic stroke

Patient	23	24	25	26	27	28	29	30
Age at presentation (years)	61	64	64	53	58	85	73	27
Gender (M/F)	М	М	М	F	М	М	М	F
Ethnicity	Black	White	White	Asian	Black	White	Asian	White
Stroke type, ob- served/implicated mech- anism	Ischaemic right mid- dle cerebral artery occlusion	Ischaemic, verte- bral-basilar artery occlusion	Ischaemic bilateral ACA-MCA and MCA-PCA cortical and deep bor- derzone infarct	Ischaemic, verte- bral-basilar artery occlusion	Ischaemic, proximal left middle cerebral artery occlusion	Ischaemic, Left pos- terior cerebral artery occlusion	Ischaemic basilar artery occlusion	Ischaemic left in- ternal cerebral ar- tery occlusion
Medical history and risk factors for stroke	Hypertension; stroke 8 years ago (left-sided weak- ness); cellulitis; in- creased body mass index	Nil	Recurrent DVTs (rivaroxaban), Conn Syndrome	Mitral valve re- placement (metal valve), atrial fibril- lation (on warfarin), heart failure, PPM in situ, hyperten- sion, type 2 diabe- tes, Grave's disease	Spondylosis	Hypertension, hy- percholesterolae- mia, atrial fibrilla- tion (on apixaban), ischaemic heart dis- ease, prostate cancer (Gleason Score 4+5)	Gastric carcinoma (resected), benign essential tremor	Nil
Days from onset of COVII	D-19 symptoms to:			•				
1) Hospital admission	-2	10	7	21	2	10	8	0
2) Onset of neurological symptoms	-2	15		22	2	10	8	0
3) ICU admission; dura- tion (days)	Did not go to ICU	Yes - 5	Yes - 13	Yes -8	Did not go to ICU	Did not go to ICU	Did not go to ICU	Yes - 7
Main reason for hospital admission	Neurological	Respiratory	Respiratory	Respiratory	Neurological	Neurological	Neurological	Neurological
Security of COVID-19 diagnosis	Definite	Definite	Definite	Definite	Probable	Definite	Definite	Probable
Chest radiograph changes	CT chest: Bilateral patchy subpleural airspace opacifica- tion in both lungs	CXR: Bilateral pulmonary infil- trates	CXR: Bilateral parenchymal air- space infiltrates, more confluent within the lower zones. Bilateral	CXR: Bilateral ground-glass changes and consol- idation	CXR: normal CT chest: Lungs clear	peripheral airspace opacities throughout both lungs, worse on right CXR: Bilat- eral	CXR: Bilateral pre- dominantly periph- eral airspace opaci- ties, most confluent at the mid-zones and lung bases	CXR: CT chest: patchy ground glass infiltrates of the left upper lobe. Other indetermi- nate infiltrates at

			shallow pleural ef- fusions					the level of both lung bases.
Severity of COVID-19 infection	Mild	Critical	Critical	Critical	Mild	Mild	Mild	Mild
Premorbid mRS	2	0	0	1	0	0	0	0
Signs and symptoms of stroke	Dysarthria, left fa- cial droop and left- sided weakness	1st event – left ataxic monoparesis of upper limb; 2nd event 7 days after (despite anticoagu- lation) - bilateral incoordination and right homonymous hemianopia	Incidental (ICU)	Acute confusion, in- coordination, re- duced conscious- ness (GCS 13/15)	Aphasia and right- sided hemiparesis	Dysarthria, right fa- cial droop	Aphasia, right facial droop and right- sided weakness	Dysphasia and left-sided hemi- paresis
Baseline NIHSS	5	2			23	6	15	15
Imaging used for diagno- sis	CT, CTA, MRI	CT, CTA, MRI	CT, CTA, MRI	CT, CTA	CT, CTA, MRI	CT, CTA	CT, CTA, MRI	CT, CTA
Risk factors for cardio- embolism ¥	No	No	Yes - AF	Yes - AF	No	Yes - AF	No	No
Significant carotid steno- sis *	No	No	No	No	No	No	No	No
Venous thromboembo- lism	Yes - PE	Yes - PE	No	No	Yes - PE	No	No	Yes -PE
Brain imaging Laboratory results within	MRI - Acute infarct in the right corpus striatum. Multiple supra- and infra-ten- torial cortical and subcortical micro- hemorrhages	MRI (1st event): acute and acute left posterior-inferior cerebellar artery territory infarction with microhaemor- rhages. 2nd event - 7 days later: bilat- eral acute posterior cerebral artery ter- ritory infarcts de- spite therapeutic anticoagulation left vertebral artery thrombus	MRI: subacute in- farcts within the deep internal bor- derzones of the cer- ebral hemispheres bilaterally, and within the left frontal white mat- ter. Background moderate small vessel disease and established cortical infarcts, in arterial borderzone territo- ries.	Non-contrast CT showed acute right parietal cortical and left cerebellar in- farct with mass ef- fect and hydroceph- alus, despite thera- peutic anticoagulan- tion.	MRI: Extensive evolving left MCA infarct with evi- dence of petechial haemorrhage and associated mass-ef- fect as described. Persistent occlusion of the left M2 MCA branches	Non-contrast CT: showed hyper- density consistent with thrombus in the left posterior cerebral artery and acute infarction in the left temporal stem and cerebral peduncle	MRI: acute infarc- tion in the right thalamus, left pons, right occipital lobe and right cerebellar hemisphere. Throm- botic material in the basilar artery and bilateral mild-to- moderate P2 seg- ment stenosis	CT: Extensive acute right middle cerebral artery and right anterior cere- bral artery territory infarction. There is local swelling and sulcal effacement with new leftward midline shift of ~4 mm.

Hb (g/L; 130-170)	126	117	70	106	147	128	159	81
MCV (fL; 80-99)	94.2	90	100.4	89.1	93.4	85	88	93.5
Lymphs (x10^9/L; 1.2- 3.65)	1.31	0.53	1.79	2.07	1.49	1.09	1.65	2.16
Neuts (x10^9/L; 2-7.5)	6.39	6.65	23.82	18.2	8.87	5.87	6.68	8.96
Plt (x10^9/L; 150-400)	408	303	294	328	128	287	632	403
CRP (mg/L; 0-5.0)	12.8	280	326.2	97.4	45	161	179.9	119
Fibrinogen (g/L; 1.5-4.0)	4.63	9.5	8.82	2.91	3.15	5.3	NR	NR
D-Dimer (µg/L; 0-550)	27190	80000	29000	7750	75320	16100	NR	NR
Prothrombin time (secs; 10-12)	10.7	11.6	12.6	34.4	12.2	11.3	14.9	11.5
APTT (secs; 25-37)	29	35	34	64.7	23	33	30	26
Cardiolipin antibody	Negative	IgM (Medium ti- tre)	IgG (low titre)	Negative	Negative	Negative	Negative	Negative
Lupus anticoagulant	Positive	Positive	NR	Positive	Positive	Negative	Negative	Positive
Beta-2 glycoprotein	Negative	IgM and IgG (low titre)	NR	Negative	Negative	Negative	IgM (low titre)	Negative
Ferritin (µg/L; 30-400)	1167	3563	1044	828	NR	NR	NR	NR
Total cholesterol (mmol/L; 2.5-5.0)	4.1	2.6	NR	2.2	5	2.9	4.1	6.2
LDL (mmol/L; 0-3.5)	2.4	1.4	NR	0.8	3.4	1.1	2.2	3.4
HDL (mmol/L; 0.9-1.5)	1	0.5	NR	0.9	0.9	0.9	0.8	1.1
Triglyceride (mmol/L; 0.4-2.3)	1.5	1.5	NR	1	1.6	1.9	2.4	3.8

Glucose (mmol/L; 3.9- 5.8)	6.2	6.9	NR	6.8	6.5	9.4	NR	NR
Management:								
Tissue Plasminogen acti- vator	No	No	No	No	No	No	Yes	No
Mechanical Throm- bectomy	No	No	No	No	No	No	No	No
Antithrombotic therapy	LMWH	LMWH	LMWH	LMWH	LMWH	aspirin seven days then switched to apixaban	aspirin five days then switched to LMWH	Aspirin 10 days then LMWH
Outcome Status	Rehabilitation unit	Rehabilitation unit	Remains static in ICU (day 31)	Died	Rehabilitation unit	Rehabilitation unit	Stroke Unit	Rehabilitation unit

Footnote: ¥ Atrial fibrillation/flutter, Severe LVEF, reduced myocardial wall motion activity, left ventricular mural thrombus - sourced from electrocardiogram, cardiac echocardiogram, and 72 hour holter monitor *significant defined by >50% narrowing. LMWH: low molecular weight heparin; mRS: modified Rankin Scale; NHISS: National Institute for Health Stroke Scale; DVT: Deep Vein Thrombosis; PPM: Permanent Pacemaker; AF: Atrial Fibrillation, MCA; Middle Cerebral Artery; ACA: Anterior Cerebral Artery; PCA: Posterior Cerebral Artery.

Table S1d. Demographic, clinical, radiological, laboratory and characteristics of individuals with possible and definite COVID-19 infection with peripheral neurological syndromes

Patient	31	32	33	34	35	36	37	38
Age at presentation (years)	61	57	63	42	60	20	38	60
Gender (M/F)	М	М	М	М	М	М	М	М
Ethnicity	White	White	Other	White	White	Black	White	White
Final neurological di- agnosis	GBS	GBS	GBS	GBS	GBS	GBS	GBS	Brachial plex- opathy
Past medical history	Cluster headache, cervical myelopathy, diabetes, arrhythmia	Hypercholesterolae- mia	Depression	Nil	Nil	Nil	Nil	Myeloma; cere- bellar stroke
Initial COVID-19 symptoms	Cough; fever; ma- laise; headache	Fever; headache; myalgia	Cough; myalgia; dyspnoea; fevers	Cough, fever dysp- noea, diarrhoea, an- osmia	Headache; ageusia; anosmia	Cough, fever, ageu- sia	Cough, diarrhoea	Cough
Days from onset of COV	VID-19 symptoms to:	·						
1) Hospital admission	15	11	31	14	0	14	23	17
2) Onset of neurologi- cal symptoms	14	6	18	13	-1	10	21	14
3) ICU admission	17	N/A	N/A	16	5	N/A	N/A	N/A
Main reason for ad- mission (neurological vs respiratory symp- toms)	Neurological	Neurological	Neurological	Neurological	Neurological	Neurological	Neurological	Neurological
Security of COVID- 19 diagnosis (definite, probable, possible)	Possible	Possible	Possible	Definite	Definite	Possible	Definite	Definite
Severity of COVID- 19 infection	Mild	Mild	Mild	Mild	Critical	Mild	Mild	Mild
Duration of ICU stay	11	NA	NA	17 days	41 days (ongoing)	NA	NA	NA

Initial neurological symptoms	Distal limb numb- ness	Distal limb weak- ness and numbness	Distal paraesthesia; limb weakness; fa- cial weakness	Distal limb numb- ness and weakness; dysphagia	Distal limb numbness and weakness	Distal limb numb- ness and weakness	Distal limb numb- ness, weakness, clumsiness	Unilateral pain- less arm weak- ness and numb- ness
Key neurological signs	Quadriparesis; are- flexia	Bilateral facial nerve palsy; bulbar weakness; distal limb weakness; are- flexia	Right facial nerve palsy; quadriparesis; areflexia; sensory ataxia	Quadriparesis; are- flexia; sensory loss	Quadriparesis; are- flexia; sensory loss; dysautonomia; facial and bulbar weakness	Mild distal weak- ness and areflexia	Mild distal weak- ness; sensory ataxia	Altered sensa- tion L C6 der- matome; weak- ness below L el- bow
Initial laboratory resul	ts at or nearest to time	of neurological sympto	om onset					
Hb (g/L; 130-170)	125	146	147	135	166	153	152	85
MCV (fL; 80-99)	95.8	82.9	90	87.7	100.1	95.1	92	107.5
Lymphs (x10^9/L; 1.2-3.65)	2.41	2.35	2.2	1.8	1.46	2.72	1.16	3.53
Neuts (x10^9/L; 2-7.5)	5.81	6.31	3.1	10.7	3.48	2.6	3.14	1.32
Plt (x10^9/L; 150-400)	252	456	199	425	256	244	180	142
CRP (mg/L; 0-5.0)		9.9	<5	6	71	<5	<5	NR
Fibrinogen (g/L; 1.5- 4.0)	NR	NR	4.1	NR	9.31	1.97	2.76	2.86
D-Dimer (µg/L; 0- 550)	NR	420	NR	NR	NR	NR	NR	9100
Ferritin (µg/L; 30-400)	NR	410	450	NR	NR	190	NR	NR
Other relevant results	Hep E serology posi- tive	NAD	NAD	NAD	NAD	NAD	NAD	NAD
CSF								
White cell count (cells/µL; 0-5)	<1	12	2	3	2	2	<1	NR
Protein (g/L; 0.13- 0.45)	0.3	1.24	0.6	0.5	0.6	0.26	0.9	NR

Glucose (mmol/L; 2.2- 4.2) (blood glucose) (mmol/L; 3.9-5.8)	NR	NR	4.0 (4.9)	NR	3.4	2.9 (4.4)	3.7 (5.4)	NR
Imaging						·		
Neuroaxis	CT brain normal	MRI brain and spine normal	MRI brain and spine normal	CT brain normal	MRI brain normal	Nerve root enhance- ment	MRI brain normal	MRI brain and spine normal
Chest	Normal	Abnormal	Normal	Normal	Normal	Normal	Normal	Midzone opaci- fication con- sistent with COVID-19
Other relevant investig	ations							
Neurophysiology	Demyelinating with severe axonal loss	Demyelinating	Not done	Demyelinating	Demyelinating	Demyelinating with axonal loss	Demyelinating	Not done
Treatment								
For neurological diag- nosis	IVIG	IVIG	IVIG	IVIG; mechanical ventilation	IVIG	IVIG	IVIG	IV methyl pred- nisolone
For COVID-19 infec- tion	Nil	Nil	Nil	Nil	Intubation and venti- lation	Nil	Nil	Nil
Outcomes		1			1	1		1
Total duration of hos- pital admission (days)	30	35	10	17	46 (ongoing)	5	7 (ongoing)	7
GBS disability score (on discharge)	4	2	2	2	5	2	2	NA

Footnote: 'NR' denotes not tested or no result. GBS: Guillian-Barre syndrome; IVIG: intravenous immunoglobulin NAD: nil abnormal detected. GBS disability score: 0 – healthy, 1 – minor symptoms or signs of neuropathy but capable of manual work/capable of running, 2 – able to walk 5m (across an open space) but incapable of manual work/running, 3 – able to walk with a stick, appliance or support (5m across an open space) 4 – bedridden or chairbound, 5 – requiring assisted ventilation (for any part of the day or night), 6 – dead.

Table S1e. Five miscellaneous and uncharacterised patients

Patient	39	40	41	42	43
Age (years)	16	17	27	40	20
Gender (M/F)	F	F	М	М	F
Ethnicity	Asian	Black	White	Asian	White
Final neurological diagnosis	Cranial nerve palsies associ- ated with pseudotumour cere- bri	Non-convulsive status epilep- ticus with widespread cortical MRI changes	Seizures; widespread micro- haemorrhages in context of crit- ical COVID-19 infection, hy- pertensive episodes and new AML on Gilteritinib	Pyogenic [Streptococcus interme- dius] CNS infection post- COVID-19	Myelopathy, cord ischaemia considered
Past medical history	Menorrhagia; normal BMI	Cornelia de Lange syndrome; epilepsy; dysmelia; hyperten- sion; visual disturbance; GORD; Nissen fundoplica- tion; gastrostomy.	Acute myeloid leukaemia (new)	Interstitial keratitis; previous wrist and ankle synovitis	None relevant
Initial COVID-19 symp- toms	Abdominal pain; diarrhoea; maculopapular rash; head- ache; sore throat; fever.	Fever; cough; dyspnoea; household contacts	Fever	Fever; cough; shortness of breath; household contacts	Cough; coryza; fatigue; myal- gia
Days from onset of COVII					
1) Hospital admission	6	3	1	14	5
2) Onset of neurological symptoms	10	26	6	14	6
3) ICU admission	N/A	5	23	14	N/A
Main reason for hospital admission (neurological vs respiratory symp- toms)	COVID-19 symptoms	Respiratory	Other	Neurological; generally unwell	Neurological
Security of COVID-19 diagnosis	Probable	Definite	Definite	Possible	Definite
Severity of COVID-19 infection	Severe	Critical	Critical	Mild	Mild
Duration of ICU stay	N/A	>50 days ongoing	46	27	N/A

Initial neurological symptoms	Diplopia; headache with fea- tures of raised ICP.	Seizures	Seizures; weakness; fatigue	Return of fever; drenching night sweats; headache; vomiting; con- fusion; low conscious level	Headache; leg weakness and numbness; constipation
Key neurological signs	Evolving cranial nerve signs; progressive bilateral failure of abduction; weakness of left orbicularis oculi; no optic disc swelling	Low conscious level; possible cortical visual impairment; in- creased tone and clonus	Mild asymmetric weakness;	Low conscious level; left-sided pyramidal weakness; brisk re- flexes	Proximal lower limb weakness; absent lower abdominal and lower limb reflexes; flexor plantars; sensory level T10
Initial laboratory results a	at or nearest to time of neurologi	cal symptom onset			
Hb (g/L; 130-170)	102	95	94	94	145
MCV (fL; 80-99)	80.1	91	94	60	84.9
Lymphs (x10^9/L; 1.2- 3.65)	1.7	1.1	0.58	1	1.6
Neuts (x10^9/L; 2-7.5)	9.3	8.6	0.74	24	2.8
Plt (x10^9/L; 150-400)	81	135	81	521	238
ESR (mm/hr; 1-20)	NR	NR	NR	103	23
CRP (mg/L; 0-5.0)	294	100	434	248	1.3
Fibrinogen (g/L; 1.5-4.0)	NR	NR	8.59	5.01	4.15
D-Dimer (normal range for assay)	626 ng/mL (0-243)	NR	22900 µg/L (0-550)	4970 µg/L (0-550)	NR
Ferritin (µg/L; 30-400)	NR	1367	6467	1592	NR
Phospholipid antibodies	NR	NR	NR	NR	Negative
Lupus anticoagulant	NR	NR	NR	NR	NR
Other relevant results	ANA and ANCA negative; thyroid function normal	Neuronal and NMDA recep- tor antibodies negative		ANA negative; Ro antibodies equivocal	ANA and ANCA negative; AQP4 and MOG antibodies negative
CSF	1	1		1	1

White cell count (cells/µL; 0-5)	Two studies performed (2nd 5 days after 1st) 1 st : <1; 2 nd : <1	3	Not done	1st: 4500 (95% lymphocytes); 2 nd : 13,920 (95% polymorphs); 3 rd : 405 (27% mononuclear cells, 73% polymorphs); 4 th : 40 (18% mononuclear cells, 82% poly- morphs); 5 th : 121 (predominantly polymorphs); 6 th 20 (75% mono- nuclear cells, 25% polymorphs)	<1
Protein (g/L; 0.13-0.45)	1st: 0.3; 2nd 0.3	0.91		1 st : 2.18; 3 rd : 1.03; 4 th : 1.42; 5 th : 0.62; 6 th : 0.54;	0.24
Glucose (mmol/L; 2.2-4.2) (blood glucose) (mmol/L; 3.9-5.8)	1st: 3.1; 2nd: 3.3	4.4		1 st : 0.7; 3 rd : NR; 4 th : 2.65; 5 th : 2.97; 6 th : 3.81	3.2
Other	Opening pressure: 1st: 39 cm H20; 2nd: 27 cm H20; CSF culture and viral PCR negative	CSF culture and viral PCR negative		Unmatched oligoclonal bands de- tected in CSF; CSF viral PCR negative, PCR for Streptococcus intermedius detected on CSF from day 5 of admission	Oligoclonal bands negative; CSF culture and viral PCR neg- ative, including SARS-CoV-2
Micro/virology	Blood and urine cultures nega- tive; Screening for other res- piratory viruses negative				
Imaging					
Neuroaxis	Dilated optic nerve sheaths and narrowed but patent trans- verse sinuses; consistent with raised intra-cranial pressure. No parenchymal changes or thrombosis of the head & neck vessels.	Extensive, symmetrical bilat- eral signal changes without restricted diffusion involving the cortices of the cerebral and cerebellar hemispheres and the thalami.	MRI brain - Extensive foci of susceptibility artefact predomi- nantly at the grey white junc- tions of cerebral hemispheres. A few foci are noted in the cere- bellum. Minimal associated sig- nal changes but no perilesional oedema or significant mass-ef- fect.	MRI brain - Bilateral ring / lami- nar enhancing cerebral and cere- bellar lesions with diffusion re- striction and partial haemorrhagic content, consistent with abscesses and apparent ventriculitis with ei- ther intraventricular pus and/or haemorrhage.	MRI brain and spine normal
Chest	CXR - bilateral infiltrates		CT chest - extensive parenchy- mal changes showing typical appearances of COVID-19 in- fection	No COVID-19 changes	
Other relevant investigation	ons				
Echocardiogram	Mild pericardial effusion, pre- served ejection fraction 55%				
Treatment	1		1	1	1
For neurological diagnosis	Therapeutic lumbar puncture; acetazolamide	Dexamethasone 4g QDS 2 days; methylprednisolone 30mg/kg 3 days, then oral prednisolone taper	Levetiracetam	Intubation and ventilation; vari- ous antimicrobials including ceftriaxone, linezolid, mero- penem, intrathecal vancomycin; ICP monitoring; right then left	Aspirin 75mg daily

				frontal EVDs; focal excision of brain abscess	
For COVID-19 infection	Colchicine for myocarditis; oxygen	Intubation and ventilation	Intubation and ventilation;	None additional	None required
Other			Chemotherapy for AML - Gilteritinib		
Outcomes					
Total duration of hospital admission (days)	18	Ongoing	Ongoing	Ongoing	5
Outcome at last follow up	Incomplete recovery; improv- ing; discharged home	No improvement; static	Incomplete recovery; improv- ing; undergoing rehabilitation	Incomplete recovery; improving	Complete recovery; discharged home

Footnote: CADASIL: Cerebral Autosomal Dominant Arteriopathy with Subcortical Infarcts and Leukoencephalopathy; TIA: transient ischaemic attack; ICU: intensive care unit; N/A: not applicable; Hb: haemoglobin; MCV: mean corpuscular volume; Lymphs: lymphocytes; Neuts: neutrophils; Plt: platelets; ESR: erythrocyte sedimentation rate; CRP: C-reactive protein; NR: no result; TSH: thyroid stimulating hormone; NMDA receptor: N-methyl-D-aspartate receptor; CSF: cerebrospinal fluid; PCR: polymerase chain reaction; CT: computed tomography; MRI: magnetic resonance imaging; CXR: chest X-ray; CTPA: CT pulmonary angiogram; ARDS: acute respiratory distress syndrome; EEG: electroencephalogram; IV: intravenous; IVMP: intravenous methylprednisolone; MGUS: monoclonal gammopathy of undetermined significance; ADEM: acute demyelinating encephalomyelitis; ANA: anti-nuclear antibodies; ANCA: anti-neutrophil cytoplasmic antibodies; ENA: extractable nuclear antigens; IgG: Immunoglobulin G; GAD: glutamic acid decarboxylase; DPPX: dipeptidyl-peptidase-like protein 6; LGI1: leucine-rich glioma-inactivated 1; CASPR2: contactin-associated protein-like 2; MOG: myelin oligodendrocyte glycoprotein; AQP4: aquaporin-4; FDG PET: fluorodeoxyglucose positron emission tomography; IVIG: intravenous immunoglobulin; HTLV 1 and 2: Human T-cell Leukaemia Virus types 1 and 2; EMG: electromyogram; ICP: intracranial pressure; EVDL external ventricular drain; Standard viral PCR – herpes simplex viruses (1 + 2), varicella zoster, enterovirus +/- adenovirus, cytomegalovirus, Epstein-Barr Virus, parechovirus depending upon NHS trust/centre; N-Gene PCR (unvalidated test) for SARS-CoV-2 testing in CSF only where specified.