

**sREMAP-CAP Amendment Tracker**

<b><u>Amendment</u></b>	<b><u>Approvals</u></b>	<b><u>Submission date</u></b>	<b><u>Approval date</u></b>	<b><u>Documents</u></b>	<b><u>Summary</u></b>
Original approvals	REC HRA MHRA	23.03.18	01.06.18		
AM01	REC HRA (linked to AM02)	18.06.18	19.07.18	REMAP-CAP European Region-Specific Appendix V2.1 20180524	1 Addition of a single recruiting site (Poole Hospital) 2 Amendment to the European Region Specific Appendix
AM02	REC HRA	13.06.18	07.08.18		1 Addition of 7 new recruiting sites (North Tees, Royal Berkshire, Sunderland Royal, Basingstoke, Royal Blackburn, QA Portsmouth, James Cook)
AM03	REC HRA	10.12.18	30.01.19	REMAP-CAP European Region specific Appendix v2.2 26.10.2018 Patient information sheet and consent form V2.1 29.11.2018 Retrospective patient information sheet and consent form V2.1 29.11.2018 Personal Legal representative information sheet and consent form V2.1 29.11.2018 Professional Legal Representative consent form V1.1 29.11.2018	1 Addition of 3 new recruiting sites (Stoke Mandeville, Peterborough City, Tunbridge Wells) 2 Change in PI at 1 site (St James' Hospital, Leeds – PI Dr E. Paramasivam) 3 Updated information sheets 4 Updated European Region Specific Appendix
AM04	REC HRA	29.07.19	03.09.19		1 Addition of 5 new recruiting sites (Milton Keynes, Darlington Memorial, North Durham, Northampton General, Royal Surrey) 2 Change in PI at one site (Royal Cornwall, New PI – Dr Mike Spivey)
AM05	REC HRA MHRA	27.11.19	09.01.20	REMAP-CAP Core Protocol V3 - FINAL_CLEAN - 10 July 2019 REMAP-CAP Macrolide Duration Domain-Specific Appendix V3 - FINAL_CLEAN - 10 July 2019 REMAP-CAP Corticosteroid Domain-Specific Appendix V3 - FINAL_CLEAN - 12 July 2019 REMAP-CAP European Region-Specific Appendix V3 - FINAL_CLEAN - 23 August 2019 REMAP-CAP Statistical Analysis Appendix V3 - FINAL_CLEAN - 24 August 2019	1 Amendments to the core protocol 2 Addition of the New domain appendices 3 Addition of an intervention to the pre-existing Corticosteroid domain

				<p>REMAP-CAP Antibiotic Domain-Specific Appendix V3 - FINAL_CLEAN - 10 July 2019</p> <p>REMAP-CAP - Antiviral Domain-Specific Appendix V1 - FINAL - 10 July 2019</p> <p>REMAP-CAP Registry Appendix v1 - UK_FINAL - 11 Sep 2019</p> <p>REMAP-CAP Protocol Summary V3 - FINAL_CLEAN - 11 Sep 2019</p> <p>UK REMAP CAP PIS ICF V2.3 29.10.2019</p> <p>UK REMAP CAP Retrospective PIS ICF V2.3 29.10.2019</p> <p>UK REMAP CAP PerLRIS ICF V2.4 29.10.2019</p>	
AM06	REC HRA	13.02.20	09.03.20	REMAP-CAP - PatC - V1.1 - CLEAN - 20200212	1 Addition of Pandemic appendix to Core
AM07	REC HRA MHRA	16.03.20	27.03.20	<p>European Region Specific Appendix v3.1 13.03.2020</p> <p>REMAP-CAP - COVID-19 Antiviral Domain-Specific Appendix V1.0 - 11 March 2020</p> <p>REMAP-CAP - COVID-19 Immune Modulation Domain-Specific Appendix V1.0 - 11 March 2020</p> <p>UK REMAP CAP PIS ICF V2.4 2020.03.16</p> <p>UK REMAP CAP Retro PIS ICF V2.4 2020.03.16</p> <p>UK REMAPCAP PerLRIS ICF v2.5 2020.03.16</p> <p>UK REMAP CAP COVID19 PIS ICF V1.0 2020.03.16</p> <p>Amended to UK REMAP CAP COVID19 PIS ICF V1.1 2020.03.19</p> <p>UK REMAP CAP COVID19 Retro PIS ICF V1.0 2020.03.16</p> <p>UK REMAPCAP COVID19 PerLRIS ICF v1.0 2020.03.16</p>	<p>1 Addition of 10 new recruiting sites (Charing Cross Hospital, Hammersmith Hospital, St Mary's Hospital, Royal Free Hospital, John Radcliffe Hospital, Addenbrookes Hospital, York Hospital, Belfast City Hospital, Royal Victoria Belfast, Mater Hospital)</p> <p>2 updated EU Region specific appendix</p> <p>3 Addition of COVID-19 Antiviral domain</p> <p>4 Addition of COVID-19 immune modulation domain</p> <p>5 updated information sheets and consent forms</p> <p>6 New COVID-19 specific information sheets and consent forms</p>
AM08	REC HRA	25.03.20	30.03.20		<p>1 Addition of co-PI at Belfast City Hospital, Royal Victoria Hospital and Mater Hospital:</p> <p>2. Addition of 21 new recruiting sites (Guy's &amp; St Thomas' Hospital, Bolton Hospital, Wythenshawe Hospital, The Cumberland Infirmary, West</p>

					Cumberland Hospital, South Tyneside Hospital, Royal United Bath, City Hospital Birmingham, Royal Brompton Hospital, Royal Marsden Hospital, Northwick Park Hospital, Ealing Hospital, Central Middlesex Hospital, Frimley Park Hospital, Wexham Hospital, Southend Hospital, Royal Gwent Hospital, King's Mill Hospital, Southampton Hospital, Salford Royal Hospital, Royal Liverpool Hospital).
AM09	REC HRA	30.03.20	09.04.20	UK REMAP CAP COVID19 PIS ICF V1.0 2020.03.28 - Scotland UK REMAP CAP COVID19 Retro PIS ICF V1.0 2020.03.28 Recovered Capacity - Scotland UK REMAPCAP COVID19 PerLRIS ICF v1.0 2020.03.28 Attorney Guardian Relative - Scotland UK REMAP-CAP ProLR ICF V1.0 2020.03.28 clean Scotland	1 Addition of 13 new recruiting sites (Bristol Royal, The Royal London, City Hospital Nottingham, Craigavon Area Hospital, Altnagelvin Hospital, NHS Nightingale Hospital, Antrim Area Hospital, Freeman Hospital, Northern General Hospital) including addition of 4 New Scottish Sites (Glasgow Royal Infirmary, Edinburgh Royal Infirmary, Queen Elizabeth Glasgow, Western General Edinburgh) 2 Change of PIs at 2 sites  3 addition of information sheets and consent forms for Scottish sites
AM10	REC HRA MHRA	06.04.20	09.04.20	REMAP-CAP - COVID-19 Antiviral Domain-Specific Appendix V2.0 - 01 April 2020_CLEAN  UK REMAP CAP COVID19 PIS ICF V1.2 2020.04.06 UK REMAP CAP COVID19 Retro PIS ICF V1.2 2020.04.06 UK REMAPCAP COVID19 PerLRIS ICF v1.2 2020.04.06	1 Addition of an additional antiviral agent, hydroxychloroquine to the COVID anti-viral domain We will evaluate this agent on its own and also in combination with kaletra.  2 We added this intervention into the information sheets and made a few minor semantic changes to make some of the terminology more lay friendly.

AM11	REC HRA MHRA	09.04.20	15.04.20	<p>REMAP-CAP - COVID-19 Immune Modulation Domain-Specific Appendix V2.0 - 07 April 2020_CLEAN</p> <p>UK REMAP CAP COVID19 PIS ICF V1.3 2020.04.09 UK REMAP CAP COVID19 Retro PIS ICF V1.3 2020.04.09 UK REMAPCAP COVID19 PerLRIS ICF v1.3 2020.04.09</p> <p>UK REMAP CAP COVID19 PIS ICF V1.1 2020.04.09 – Scotland UK REMAP CAP COVID19 Retro PIS ICF V1.1 2020.04.09 Recovered Capacity – Scotland UK REMAPCAP COVID19 PerLRIS ICF v1.1 2020.04.09 Attorney Guardian Relative – Scotland UK REMAP CAP COVID19 PerLRIS ICF v1.2 2020.04.09_NI UK REMAP CAP COVID19 PIS ICF V1.2 2020.04.09_NI UK REMAP CAP COVID19 Retro PIS ICF V1.2 2020.04.09_NI</p>	<p>1 changes to the Immune modulation DSA-2 additional immune modulatory agent, Tocilizumab and Sarilumab to the COVID immune modulation domain of the REMAP-CAP trial.</p> <p>2 We added these interventions into the England, Northern Ireland and Scotland information sheets and made a few minor changes to make some of the terminology more lay friendly.</p>
AM12	HRA	15.04.20	15.04.20	Substantial amendment letter generated and submitted by the HRA	<p>All UK NHS organisations that have ICU's / Critical care facilities. This includes all NHS ICU units in England and in the devolved states so that they may enrol and randomise patients with acute COVID-19 infection into the REMAP-CAP study.</p> <p>All Principal Investigators at these organisations, as deemed appropriate by the sponsor. This should be taken to include cases where the PI is changed at a previously approved NHS site.</p>
AM13	REC HRA	19.04.20	22.04.20	<p>REMAP-CAP COVID-19 Immunoglobulin Therapy Domain-Specific Appendix V1- 19 April 2020</p> <p>UK REMAP CAP COVID19 PIS ICF V1.4 2020.04.19 UK REMAP CAP COVID19 Retro PIS ICF V1.4 2020.04.19 UK REMAPCAP COVID19 PerLRIS ICF v1.4 2020.04.19</p> <p>UK REMAP CAP COVID19 PerLRIS ICF v1.3 2020.04.19_NI UK REMAP CAP COVID19 PIS ICF V1.3 2020.04.19_NI</p>	<p>1 Addition of the COVID-19 Immunoglobulin Domain specific appendix v1 19.04.20.</p> <p>2 We added these interventions into the information sheets and made a few minor semantic changes to make some of the terminology more lay friendly.</p>

				<p>UK REMAP CAP COVID19 Retro PIS ICF V1.3 2020.04.19_NI</p> <p>UK REMAP CAP COVID19 PIS ICF V1.2 2020.04.19 - Scotland</p> <p>UK REMAPCAP COVID19 PerLRIS ICF v1.2 2020.04.19 Attorney Guardian Relative - Scotland</p> <p>UK REMAP CAP COVID19 Retro PIS ICF V1.2 2020.04.19 Recovered Capacity - Scotland</p>	
AM14	REC HRA MHRA	22.04.20	27.04.20	<p>REMAP-CAP COVID-19 Therapeutic Anticoagulation Domain-Specific Appendix V1- 20 April 2020</p> <p>UK REMAP-CAP ProLR ICF V1.2 2020.04.20</p> <p>UK REMAP CAP COVID19 PIS ICF V1.5 2020.04.22</p> <p>UK REMAP CAP COVID19 Retro PIS ICF V1.5 2020.04.22</p> <p>UK REMAPCAP COVID19 PerLRIS ICF v1.5 2020.04.22</p> <p>UK REMAP CAP COVID19 PerLRIS ICF v1.4 2020.04.22_NI</p> <p>UK REMAP CAP COVID19 PIS ICF V1.4 2020.04.22_NI</p> <p>UK REMAP CAP COVID19 Retro PIS ICF V1.4 2020.04.22_NI</p> <p>UK REMAP CAP COVID19 PIS ICF V1.3 2020.04.22 - Scotland</p> <p>UK REMAPCAP COVID19 PerLRIS ICF v1.3 2020.04.22 Attorney Guardian Relative - Scotland</p> <p>UK REMAP CAP COVID19 Retro PIS ICF V1.3 2020.04.22 Recovered Capacity - Scotland</p>	<p>1 Addition of the Therapeutic Anticoagulation Domain specific appendix v1 20.04.20.</p> <p>2 We added this intervention into the information sheets and made a few minor semantic changes to make some of the terminology more lay friendly.</p>

AM15	REC HRA MHRA	06.05.20	12.05.20		<p>Doses of Anakinra approved as per AMO11, but not updated on the CTA.</p> <p><b>New dosing from Immune modulation domain v2:</b>  Anakinra will be administered as an intravenous bolus injection via a central or peripheral line. A loading dose of 300 mg will be administered, followed by maintenance doses of 100 mg of anakinra administered every 6 hours. In patients with creatinine clearance of less than 30 ml/min or receiving renal replacement therapy, anakinra will be dosed every 12 hours.  Duration of therapy  Anakinra will be administered four times daily until the patient has been breathing without receiving invasive mechanical ventilation for more than 24 hours or for 14 days in patients who continue to receive invasive mechanical ventilation. For patients not receiving invasive mechanical ventilation, the drug will stop on ICU discharge or after 14 days, whichever occurs first.</p> <p>Correction to an error in our original MHRA CTA application regarding specific populations of patient included in this study. We had ticked excluding women of child bearing age (using and not using contraception) , but as discussed with the MHRA at that time, these patients would be included for some IMPs but not others. In light of the pandemic we feel this population of patients as well as pregnant women and breastfeeding</p>
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					women, would benefit from this study and would like to ensure they are able to be included. Pregnant and breastfeeding women are excluded from certain domains and if treating clinicians feel it would not be in their best interest to enter the study, they can be excluded on an individual basis
AM016	MHRA	20.05.20	22.05.20		<p>Addition of 2 IMP products to the CTA</p> <ol style="list-style-type: none"> <li>1. The addition of unlicensed Sarilumab. The manufacturer (Sanofi Montpellier) will be providing the study with both licenced and unlicensed product for use in REMAP-CAP. The unlicensed product will be provided with clinical trial labels (submitted to MHRA), GMP certification and QP declaration. Sarilumab will be supplied to trial sites through PHE authorised distribution sites</li> <li>2. The addition of Kaletra 80mg/20mg Oral Solution.</li> </ol>
AM017	REC HRA	23.05.20	26.05.20		<p>MHRA requested a pause to hydroxychloroquine allocation within the REMAP-CAP study.</p> <p>The pause to this intervention was part of an urgent safety measure.</p> <p>For patients who are currently being treated with hydroxychloroquine, an</p>

					individual risk assessment by the treating clinician should be made as to whether to continue or stop the treatment as per protocol.
AM018	MHRA	01.06.20	03.06.20	MHRA CTA	Addition of releasing site information to section D9 of the MHRA CTA for unlicensed Sarilumab and the Kaletra oral solution
AM019	REC HRA MHRA	05.06.20	11.06.20	REMAP-CAP COVID-19 Immunoglobulin Therapy Domain-Specific Appendix V1.1- 01 June 2020 Summary of changes to Pandemic Appendix to Core Protocol V1.0 - 24 May 2020 REMAP-CAP - PATC - V2.0 - 18 May 2020_CLEAN	Changes to Pandemic Appendix to Core – Alignment of REMAP-CAP and REMAP-COVID in readiness for pre-ICU randomisation for certain domains, and the introduction of moderate and severe patients.  Changes to Immunoglobulin domain.
AM020	Rec HRA MHRA	10.07.20	15.07.20	<b>Updates to the Therapeutic Anticoagulation Domain Specific Appendix:</b> REMAP-CAP - Therapeutic Anticoagulation - V2 - 24 June 2020_TC REMAP-CAP - Therapeutic Anticoagulation - V2 - 24 June 2020_CLEAN Summary of changes to Therapeutic Anticoagulation Domain-Specific Appendix V1.0 - 24 June 2020_FINAL The main change to this domain is to include patients with COVID-19 before they are admitted to ICU. We have therefore made changes to the participant information sheets to include these patients. We have reformatted the documentation to be more modular (for use in the general hospital ward and in the intensive care) and simplified and clarified the wording in various places. This reformatting has also been completed for the non-pandemic IS_CF documentation.  <b>The addition of 2 new Domain specific Appendices:</b> REMAP-CAP Vitamin C Domain Specific Appendix V2 - 08 June 2020 for patients with COVID-19 and community acquired pneumonia ( <b>not yet active</b> ) REMAP-CAP Simvastatin Domain-Specific Appendix - V1.0 - 07 July 2020 for patients with COVID-19 ( <b>not yet active</b> )	



				<p><b>Updated information sheets :</b></p> <p><b>Pandemic Patients</b></p> <p><b>England</b>  UK REMAP CAP COVID19 PIS ICF V1.6 2020.10.07  UK REMAP CAP COVID19 Retro PIS ICF V1.6 2020.07.10  UK REMAPCAP COVID19 PerLRIS ICF v1.6 2020.07.10  UK REMAP-CAP ProLR ICF V1.3 2020.07.10 TC</p> <p><b>Northern ireland</b>  UK REMAP CAP COVID19 PerLRIS ICF v1.5 2020.07.10_NI  UK REMAP CAP COVID19 PIS ICF V1.5 2020.07.10_NI  UK REMAP CAP COVID19 Retro PIS ICF V1.5 2020.07.10_NI  UK REMAP-CAP ProLR ICF V1.3 2020.07.10 TC</p> <p><b>Scotland</b>  UK REMAP CAP COVID19 PIS ICF V1.4 2020.07.10 – Scotland  UK REMAP CAP COVID19 Retro PIS ICF V1.4 2020.07.10 Recovered  Capacity – Scotland  UK REMAPCAP COVID19 PerLR IS ICF v1.4 2020.07.10 Attorney  Guardian Relative – Scotland  UK REMAP-CAP ProLR ICF V1.3 2020.07.10 TC</p> <p><b>Non Pandemic Patients</b></p> <p><b>England</b>  UK REMAP CAP PIS ICF V2.5 2020.07.10  UK REMAP CAP Retro PIS ICF V2.5 2020.07.10  UK REMAPCAP PerLR IS ICF v2.6 2020.07.10  UK REMAP-CAP ProLR ICF V1.3 2020.07.10 TC</p> <p><b>Northern Ireland</b>  UK REMAP CAP PerLR IS ICF v2.6 2020.07.10_NI  UK REMAP CAP PIS ICF V2.5 2020.07.10_NI  UK REMAP CAP Retro PIS ICF V2.5 2020.07.10_NI</p>	
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				<p>UK REMAP-CAP ProLR ICF V1.3 2020.07.10 TC</p> <p><b>Scotland</b>  UK REMAP CAP PerLR IS ICF v2.6 2020.07.10_ Attorney Guardian Relative – Scotland  UK REMAP CAP PIS ICF V2.5 2020.07.10_Scotland  UK REMAP CAP Retro PIS ICF V2.5 2020.07.10_ Recovered Capacity – Scotland  UK REMAP-CAP ProLR ICF V1.3 2020.07.10 TC</p> <p>Dexamethasone is now considered a standard of care for most patients with COVID-19 requiring oxygen or other ventilatory support. For this reason, we have halted randomisation into the corticosteroid domain in REMAP-CAP <u>for patients known or suspected to have COVID-19</u>. This domain has been removed from the COVID-19 participant information sheets. The corticosteroid domain remains active for non-COVID-19 community acquired pneumonia.</p> <p>Hydroxychloroquine interventions have been halted in the REMAP-CAP COVID-19 Antiviral Domain. These interventions have been removed from the participant information sheets.</p>	
AM021	REC HRA MHRA	23.07.20	03.08.20	<p>REMAP-CAP Simvastatin domain specific appendix v1.1 23.07.2020</p>	<p>Update to Simvastatin DSA to add an exclusion criteria - <b><i>“Women who are known or suspected to be pregnant or breastfeeding will be excluded from the simvastatin intervention”</i></b></p>
AM022	REC HRA MHRA	1.10.201	12.10.20	<p><b>Antiplatelet Domain specific Appendix:</b>  REMAP-CAP – Antiplatelet Domain v1.0 24/08/2020 CLEAN</p> <p>UK REMAP CAP COVID19 PerLRIS ICF v1.6 2020.10.01 NI</p> <p>UK REMAP CAP COVID19 PIS ICF V1.6 2020.10.01 NI</p>	<p><b>This involves the addition of an Antiplatelet Domain specific Appendix:</b>  REMAP-CAP – Antiplatelet Domain v1.0 24/08/2020 CLEAN  Changes to IS_CF for COVID-19 suspected /confirmed patients now includes a section for randomising to</p>

				<p>UK REMAP CAP COVID19 Retro PIS ICF V1.6 2020.10.01 NI</p> <p>UK REMAP CAP COVID19 PIS ICF V1.7 2020.10.01</p> <p>UK REMAPCAP COVID19 PerLRIS ICF v1.7 2020.10.01</p> <p>UK REMAPCAP COVID19 Retro PIS ICF v1.7 2020.10</p> <p>UK REMAP CAP COVID19 PerLR ICF V1.5 2020.10.01 Scotland</p> <p>UK REMAP CAP COVID19 PIS ICF V1.5 2020.10.01 Scotland</p> <p>UK REMAP CAP COVID19 Retro PIS ICF V1.5 2020.10.01 Scotland</p> <p>We have also listed the new IMPs for the Antiplatelet Domain (SmPCs attached) :</p> <p>Aspirin Prasugrel Clopidogrel Ticagrelor</p>	<p>the antiplatelet domain for pre-ICU and ICU patients. We have also added space for witness consent in the event that the patient or the Personal Legal representative is not able to physically sign the consent form. I have attached and submitted both the TC and clean versions of the documents</p>
AM023	HRA REC	13.11.20	17.11.20	<p>Brief REMAP-CAP Patient information summary v1 13.11.20</p> <p>Brief REMAP-CAP ProLR information summary v2 13.11.20</p> <p>REMAP-CAP COVID poster v1 10.11.20</p>	<p>This amendment includes the addition of:</p> <p>A simple one page lay summary of the REMAP-CAP study for patients with COVID who have capacity but are unable to read /understand the full patient information sheets at the time of randomisation. Recruiting sites have indicated this would be an invaluable addition in the recruitment process, as some patients have sufficient capacity to say they wish to take part in the study but are unable to absorb all the information provided in the participant information sheets when they are acutely unwell in an</p>

					<p>emergency setting. This document will also allow patients to better understand the key principles of the study in advance of recruitment and confirm that they will review and sign the full information sheet and consent form at a later stage when they have sufficient capacity.</p> <p>A simple one page REMAP-CAP study summary as supportive information for Professional Legal Representatives in the event they require some simple information prior to signing the ProLR consent form.</p> <p>A simple REMAP-CAP COVID-19 waiting room poster that can be made available to patients and Personal Legal Representatives to increase awareness of the study for ICU and ward based recruitment.</p>
AM023	HRA REC	13.11.20	17.11.20, reissued ( with correction) 27.11.20)	<p>Brief REMAP-CAP Patient information summary v1 13.11.20</p> <p>Brief REMAP-CAP ProLR information summary v1 13.11.20</p> <p>Corrected to Brief REMAP-CAP ProLR information summary v2 13.11.20</p> <p>REMAP-CAP COVID poster v1 10.11.20</p>	<p>Brief REMAP-CAP Patient information summary v1 13.11.20</p> <p>Brief REMAP-CAP ProLR information summary v2 13.11.20</p> <p>REMAP-CAP COVID poster v1 10.11.20</p>
AM024	MHRA	16.02.21	04.03.21	MHRA CTA	Addition of clinical trial stock of Vitamin C
AM025	REC HRA MHRA	09.03.21	17.03.21	<p>Brief REMAP-CAP Patient information summary v2 04.03.21</p> <p>Brief REMAP-CAP ProLR information summary v3 04.03.21</p> <p>UK REMAP CAP COVID19 PerLRIS ICF v1.7 02.03.21 NI</p> <p>UK REMAP CAP COVID19 PIS ICF V1.7 02.03.21 NI</p> <p>UK REMAP CAP COVID19 Retro PIS ICF V1.7 02.03.21 NI</p> <p>UK REMAP CAP COVID19 PIS ICF V1.8 02.03.21</p>	<p>Addition of ACE2 RAS domain</p> <p>Addition of new interventions for anticoagulation domain</p> <p>Increase of patient recruitment number from 1000 to 6000</p> <p>Update all full information sheets and consent forms</p> <p>Update lay summary information sheets</p>

				<p>UK REMAPCAP COVID19 PerLRIS ICF v1.8 02.03.21</p> <p>UK REMAPCAP COVID19 Retro PIS ICF v1.8 02.03.21</p> <p>UK REMAP CAP COVID19 PerLR ICF V1.6 02.03.21 Scotland</p> <p>UK REMAP CAP COVID19 PIS ICF V1.6 02.03.21 Scotland</p> <p>UK REMAP CAP COVID19 Retro PIS ICF V1.6 02.03.21 Scotland</p>	
AM026	REC HRA MHRA	03.06.21	REC and HRA approval 29.07.21 (AM026 and AM027 approved together) MHRA rejected this amendment and the IMP specific documentation was resubmitted as part of AM027	<p>RECOVER funding agreement</p> <p>UK REMAP CAP COVID19 PerLRIS ICF v1.8 01.06.21 NI</p> <p>UK REMAP CAP COVID19 PIS ICF v1.8 01.06.21 NI</p> <p>UK REMAP CAP COVID19 Retro PIS ICF v1.8 01.06.21 NI</p> <p>UK REMAP CAP COVID19 PIS ICF v1.9 01.06.21</p> <p>UK REMAPCAP COVID19 PerLRIS ICF v1.9 01.06.21</p> <p>UK REMAPCAP COVID19 Retro PIS ICF v1.9 01.06.21</p> <p>UK REMAP CAP COVID19 PerLR ICF v1.7 01.06.21 Scotland</p> <p>UK REMAP CAP COVID19 PIS ICF v1.7 01.06.21 Scotland</p> <p>UK REMAP CAP COVID19 Retro PIS ICF v1.7 01.06.21 Scotland</p>	<p>Addition of the RECOVER funding agreement (since the PREPARE agreement funding ended); extension in recruitment period till 13.02.22</p> <p>Closure of the antiplatelet domain.</p> <p>The addition of the DMX-200 intervention (ACE2/RAS domain) and their addition to the Participant information sheets</p>
AM027	REC HRA MHRA	30.06.21	29.07.21	<p>Addition of the DMX-200 intervention to the ACE2/RAS Domain</p> <p>DIL2501 REMAP CAP label Proof english &amp; dutch 20210505</p> <p>DMX-200 ARDS IB v4 SIGNED</p> <p>DMX-200 ARDS IB v4.0 track changes</p>	

				<p>DMX-200 COVID-19 IMPD v2.0 April 2021  DMX-200_REMAP_CAP_label_proof_Part_2_sign_of 20210506  Grant Agreement-101003589-RECoVER_FE  IRASEudractExport  MhraProductsForm (AM026)  MIA_Fisher_DE_BW_01_MIA_2021_0022_DE+ EN  QP Declaration REMAP CAP_27May2021  Site GMP Certificate FDA EU and Registration Database Reference  2020 Signed</p>	
AM028	REC HRA MHRA	21.08.21	06.10.21	<p>Cysteamine_Nylexa IMPD_V1.0_18 Aug 2021  Nylexa clinical label_Version4.0_13 Jul 2021  REMAP-CAP - Domain Specific Appendix - Cysteamine - V1.0 - 21  June 2021  REMAP-CAP pharmacy guide - Version 3 - 18 August 2021_CLEAN  IRASEudractExport  MhraProductsForm AM028  Updated COVID-19 participant information sheets</p> <p>Brief REMAP-CAP Patient information summary v3 16.08.21  Brief REMAP-CAP ProLR information summary v4 16.08.21</p> <p>UK REMAP CAP COVID19 PerLRIS ICF v1.9 16.08.21 NI</p> <p>UK REMAP CAP COVID19 PIS ICF v1.9 16.08.21 NI</p> <p>UK REMAP CAP COVID19 Retro PIS ICF v1.9 16.08.21 NI</p> <p>UK REMAP CAP COVID19 PIS ICF v1.10 16.08.21  UK REMAPCAP COVID19 PerLRIS ICF ICF v1.10 16.08.21</p> <p>UK REMAPCAP COVID19 Retro PIS ICF v1.10 16.08.21</p>	

				<p>UK REMAP CAP COVID19 PerLR ICF ICF v1.8 16.08.21 Scotland</p> <p>UK REMAP CAP COVID19 PIS ICF v1.8 16.08.21 Scotland  UK REMAP CAP COVID19 Retro PIS ICF v1.8 16.08.21 Scotland</p> <p>Updated Non-pandemic participant information sheets:  <b>England</b>  UK REMAP CAP PIS ICF V2.6 16.08.21  UK REMAP CAP Retro PIS ICF V2.6 16.08.21  UK REMAPCAP PerLR IS ICF v2.7 16.08.21</p> <p><b>Northern Ireland</b>  UK REMAP CAP PerLR IS ICF v2.7 16.08.21_NI  UK REMAP CAP PIS ICF V2.6 16.08.21_NI  UK REMAP CAP Retro PIS ICF V2.6 16.08.21_NI</p> <p><b>Scotland</b>  UK REMAP CAP PerLR IS ICF v2.7 16.08.21_ Attorney Guardian  Relative – Scotland  UK REMAP CAP PIS ICF V2.6 16.08.21_Scotland  UK REMAP CAP Retro PIS ICF V2.6 16.08.21_ Recovered Capacity –  Scotland</p>	
AM029	HRA MHRA	22.09.21	06.10.21	<p>We added 1 IMP product to the MHRA CTA  Cysteamine Bitartrate</p> <p>For AM029 I have submitted the below documents to both the  MHRA and the HRA Portals:  Amendment_Tool_v1_5_25Mar21 REMAP-CAP AM029 22.09.21  Locked  Cysteamine_Nylexa IMPD_V1.0_18 Aug 2021  Nylexa clincial label_Version4.0_13 Jul 2021  REMAP-CAP - Domain Specific Appendix - Cysteamine - V1.0 - 21  June 2021  REMAP-CAP pharmacy guide - Version 3 - 18 August 2021_CLEAN  MhraProductsForm_AM028 200821</p>	<p>Much of this documentation was  submitted as part of AM028, however  the MHRA required additional  information be provided within the IB  and so provided a notice of non-  acceptance for AM028. The IMP  documentation, supplied as part of  AM028, plus the updated IB (track  changed and clean) is now being  resubmitted as AM029 for review and  approval by the MHRA and HRA. All  patient facing documentation remains  as part of AM028 for review and  approval by the REC / HRA.</p>

				IRASEudractExport Cysteamine_Nylexa IB_Version1.1_22 Sep 2021_Track changed Cysteamine_Nylexa IB_Version1.1_22 Sep 2021	
AM030	REC HRA ( Not approved) MHRA (Not approved)	30.11.21	Not approved, Amended and Resubmitted as AM032	<p><b>Documents submitted as part of Substantial amendment AM030:</b></p> <p>REMAP-CAP - ACE2 RAS DSA - V2 - 14 Oct 2021  REMAP-CAP - ACE2 RAS DSA - V2 - 14 Oct 2021_TC  Summary of changes to ACE2 RAS Domain-Specific Appendix V1.0 - 16 Nov 2021  REMAP-CAP - Monoclonal Antibody DSA V1 22 November 2021  REMAP-CAP COVID-19 Immunoglobulin Therapy Domain-Specific Appendix V3 - 24 November 2021  REMAP-CAP COVID-19 Immunoglobulin Therapy Domain-Specific Appendix V3 - 24 November 2021_TC from V1.01  Amendment_Tool_AM30_20211201</p> <p>revised-gb-spc-ronapreve-clean-120mg-ml12aug2021docx  TRV027 Vial Carton Label version3 REMAP-CAP 19Jul21  TRV027 IMPD V4.0 26August2021  TRV027 IMPD V4 Signature page  TRV027 IB Ed 7 30Nov2021</p> <p>Brief REMAPCAP ProLR_IS summary v5 261121 TC  Brief REMAPCAP ProLR_IS summary v5 261121 CLEAN  UK REMAP CAP COVID19 PerLR ICF V1.9 2021.11.26 Scotland TC  UK REMAP CAP COVID19 PerLR ICF V1.9 2021.11.26 Scotland CLEAN  UK REMAP CAP COVID19 PerLRIS ICF v1.10 2021.11.26 NI TC  UK REMAP CAP COVID19 PerLRIS ICF v1.10 2021.11.26 NI CLEAN  UK REMAP CAP COVID19 PerLRIS ICF v1.11 2021.11.26 TC  UK REMAP CAP COVID19 PerLRIS ICF v1.11 2021.11.26 CLEAN  UK REMAP CAP COVID19 PIS ICF V1.9 2021.11.26 Scotland TC  UK REMAP CAP COVID19 PIS ICF V1.9 2021.11.26 Scotland CLEAN  UK REMAP CAP COVID19 PIS ICF V1.10 2021.11.26 NI TC  UK REMAP CAP COVID19 PIS ICF V1.10 2021.11.26 NI CLEAN  UK REMAP CAP COVID19 PIS v1.11 TC  UK REMAP CAP COVID19 PIS v1.11 CLEAN  UK REMAP CAP COVID19 Retro PIS ICF V1.9 2021.11.26 Scotland TC  UK REMAP CAP COVID19 Retro PIS ICF V1.9 2021.11.26 Scotland CLEAN  UK REMAP CAP COVID19 Retro PIS ICF V1.10 2021.11.26 NI TC</p>	<p><b>This involves:</b></p> <p><b>The addition of a new intervention, TRV027 into the ACE2 RAS domain</b>  <b>The reopening of the immunoglobulin domain with new interventions</b>  <b>The addition of the Monoclonal Antibody Domain</b>  <b>I also made the below changes to the MHRA CTA.</b>  TRV027  casirivimab / imdevimab (Ronapreve)</p>



				UK REMAP CAP COVID19 Retro PIS ICF v1.10 2021.11.26 NI CLEAN UK REMAP CAP COVID19 Retro PIS ICF v1.11 2021.11.26 TC UK REMAP CAP COVID19 Retro PIS ICF v1.11 2021.11.26 CLEAN	
AM031	MHRA	17.12.21	20.12.21	<ol style="list-style-type: none"> <li>1. IMPD v2.0 dated 13 Dec 2021_track changed</li> <li>2. IMPD v2.0 dated 13 Dec 2021_Final</li> <li>3. Cysteamine label v5.0 dated 20 Aug 2021 – Expiry date extended up to Jun 2022</li> </ol>	<p>The main changes in the revised IMPD include:</p> <ol style="list-style-type: none"> <li>1. Change of AMRI name to Curia (Scotland) Limited – page 101</li> <li>2. Inclusion of 18m stability report for non-GMP batch and 6month stability report for GMP batch : Section 2.1.P.&amp; Stability – p 171 onwards. This data supports extension of expiry date to Jun 2022</li> <li>3. Shelf-life extension plan – page 177-178</li> </ol>
AM032	REC HRA MHRA	17.12.21	11.01.22	<p><b><u>Documents submitted as part of Substantial amendment AM030:</u></b>  REMAP-CAP - Monoclonal Antibody DSA V1 22 November 2021  REMAP-CAP COVID-19 Immunoglobulin Therapy Domain-Specific Appendix V3 - 29 November 2021 CLEAN  REMAP-CAP COVID-19 Immunoglobulin Therapy Domain-Specific Appendix V3 - 24 November 2021_TC from V1.01  REMAP-CAP Amendment_Tool_AM32_20211217</p> <p>revised-gb-spc-ronapreve-clean-120mg-ml12aug2021docx</p> <p>Brief REMAPCAP ProLR_IS summary v5 261121 TC  Brief REMAPCAP ProLR_IS summary v5 261121 CLEAN  UK REMAP CAP COVID19 PerLR ICF V1.9 2021.11.26 Scotland TC  UK REMAP CAP COVID19 PerLR ICF V1.9 2021.11.26 Scotland CLEAN  UK REMAP CAP COVID19 PerLRIS ICF v1.10 2021.11.26 NI TC  UK REMAP CAP COVID19 PerLRIS ICF v1.10 2021.11.26 NI CLEAN  UK REMAP CAP COVID19 PerLRIS ICF v1.11 2021.11.26 TC  UK REMAP CAP COVID19 PerLRIS ICF v1.11 2021.11.26 CLEAN  UK REMAP CAP COVID19 PIS ICF V1.9 2021.11.26 Scotland TC  UK REMAP CAP COVID19 PIS ICF V1.9 2021.11.26 Scotland CLEAN</p>	<p><b>The reopening of the immunoglobulin domain with new interventions</b>  <b>The addition of the Monoclonal Antibody Domain</b>  <b>The below changes to the MHRA CTA.</b>  casirivimab / imdevimab (Ronapreve)</p>

				<p>UK REMAP CAP COVID19 PIS ICF v1.10 2021.11.26 NI TC  UK REMAP CAP COVID19 PIS ICF v1.10 2021.11.26 NI CLEAN  UK REMAP CAP COVID19 PIS v1.11 TC  UK REMAP CAP COVID19 PIS v1.11 CLEAN  UK REMAP CAP COVID19 Retro PIS ICF v1.9 2021.11.26 Scotland TC  UK REMAP CAP COVID19 Retro PIS ICF v1.9 2021.11.26 Scotland CLEAN  UK REMAP CAP COVID19 Retro PIS ICF v1.10 2021.11.26 NI TC  UK REMAP CAP COVID19 Retro PIS ICF v1.10 2021.11.26 NI CLEAN  UK REMAP CAP COVID19 Retro PIS ICF v1.11 2021.11.26 TC  UK REMAP CAP COVID19 Retro PIS ICF v1.11 2021.11.26 CLEAN</p>	
AM033	MHRA	17.02.22		<p>MhraProductsForm_REMAP-CAP AM033 170222  MIA (IMP) 18693 v 41  REMAP-CAP DMX-200 MIA (IMP) Certificate of GMP Compliance 03Nov2021  REMAP-CAP AM033 Amendment_Tool_v1_6_LOCKED</p>	<p>Addition of UK QP oversight of DMX-200 by Fisher Clinical services UK LTD, Langhurstwood road, Horsham, RH12 4QD, UK.  Submission of associated MIA IMP  Submission of Certificate of GMP compliance for Fisher Clinical services, Horsham.</p>
AM034	REC/HRA (Information) MHRA (Review)	03.03.22	13.04.22	<p>MhraProductsForm_REMAP-CAP AM034 020322  Ascorbic Acid Smpc  REMAP-CAP AM034 Amendment_Tool_v1_6_LOCKED</p>	<p>Addition of releasing site information to section D9 of the MHRA CTA for licensed ascorbic acid  <b>This involves changes to the MHRA CTA for the Ascorbic Acid stock for use within REMAP-CAP</b>  <b>Stock: 50ml vials of concentrate for solution for injection/infusion contains: 7,5g ascorbic acid</b>  <b>Addition of UK Releasing site: Alliance Healthcare; Unit 1, High View Road Ind Units, South Normanton, Derbyshire; DE55 2DT</b>  <b>Marketing Authorisation holder: Pascoe pharmazeutische Präparate GmbH</b>  <b>UK MA number: PL 14369/0009</b>  <b>Submission of associated SmpC</b></p>

					<b>We are also formally submitting an amendment for the Pause of the ACE2/RAS domain within REMAP-CAP</b>
AM035	REC /HRA /MHRA	12.04.22	12.05.22	MhraProductsForm_REMAP-CAP AM035 110422 Cysteamine_Nylexa IMPD_V3.0_08 Apr 2022_Final Cysteamine_Nylexa IMPD_V3.0_08 Apr 2022_track changed REMAP-CAP AM035 Amendment_Tool_v1_6_LOCKED GB_REMAP-CAP_SubjectInsurance_Allianz_Endorsement 2_20210715 GB_REMAP-CAP_SubjectInsurance_Allianz_Endorsement 1_20210125 Subject Insurance Certificate	Addition of releasing site information to section D9 of the MHRA CTA for cysteamine. <b>This involves changes to the MHRA CTA for the Cysteamine stock for use within REMAP-CAP</b> <b>Addition of UK Releasing site: Catalent CTS (Edinburgh) Limited</b> ,Unit 1, Inchwood Park, Bathgate,West Lothian, EH48 2FY, United Kingdom <b>Marketing Authorisation holder: Catalent CTS (Edinburgh) Limited</b> , <b>MIA (IMP) number: 25274</b> <b>Submission of associated IMPD V3 08.04.22 (tracked changes and final)</b> <b>Update in clinical trial insurance to cover till 01.06.2023</b>
AM036	REC / HRA / MHRA	31.05.2022	26.06.22	REMAP-CAP AM036 Amendment_Tool_v1_6_LOCKED REMAP-CAP_Protocol_EuropeanRSA_V3.3_20200408_eng REMAP-CAP_Protocol_EuropeanRSA_V3.3_20200408_eng_TC REMAP-CAP_Protocol_EuropeanRSA_V3.4_20200420_eng REMAP-CAP_Protocol_EuropeanRSA_V3.4_20200420_eng_TC REMAP-CAP_Protocol_EuropeanRSA_V3.5_20200503_eng REMAP-CAP_Protocol_EuropeanRSA_V3.5_20200503_eng_TC REMAP-CAP_European RSA_ChangeHistory_20210910_eng_2 Cysteamine_Nylexa IMPD_V4.0_25 May 2022_Final Cysteamine_Nylexa IMPD_V4.0_25 May 2022_track changed RECOVER Amendment-Funding to 2023	Updates to the Region-Specific Appendix, Significant changes to the IMPD, Extension of funding for the study until December 2023 and pausing the anticoagulation domain.
AM037	REC HRA MHRA	14.11.2022	15.12.2022	REMAP-CAP AM037 Amendment_Tool_v1_6_LOCKED REMAP-CAP - SoC to CORE protocol V3.1 - 11Nov2022 REMAP-CAP Core Protocol V3.1 (UK) - 9 Nov 2022_TC REMAP-CAP Core Protocol V3.1 (UK) - 9 Nov 2022_CLEAN REMAP-CAP - SOC to Corticosteroid DSA V2 - 11Nov2022	<ul style="list-style-type: none"> <li>Submission of the Core Protocol (V3.1 tracked changes and final) – study now includes adults and children on the wards and ICU</li> </ul>

				<p>REMAP-CAP Corticosteroid Domain-Specific Appendix V5 (UK)_20221109_TC from V3</p> <p>REMAP-CAP Corticosteroid Domain-Specific Appendix V5 (UK)_20221109_CLEAN</p> <p>REMAP-CPA - SoC to Influenza Antiviral Domain-Specific Appendix - V1_11Nov22</p> <p>REMAP-CAP - Influenza Antiviral Domain-Specific Appendix - V2 (UK) - 01 Nov 2022_TC</p> <p>REMAP-CAP - Influenza Antiviral Domain-Specific Appendix - V2 (UK) - 01 Nov 2022_CLEAN</p> <p>Dexamethasone 3.3 mg/ml Solution for Injection (vial) - Summary of Product Characteristics (SmPC)</p> <p>Xofluza 20 mg &amp; 40 mg film coated tablets - Summary of Product Characteristics (SmPC)</p> <p>Tamiflu 30 mg Hard Capsules - Summary of Product Characteristics (SmPC)</p> <p>REMAP-CAP_Tel v1.0 25OCT22</p> <p>REMAP-CAP_Privacy Notice v1.0 25OCT22</p> <p>REMAP-CAP_PIS_ICF v1.12 2022.10.25_tracked</p> <p>REMAP-CAP_PIS_ICF v1.12 25OCT22</p> <p>REMAP-CAP_PerLIS_ICF v1.12 2022.10.25_tracked</p> <p>REMAP-CAP_PerLIS_ICF v1.12 25OCT22</p> <p>REMAP-CAP_PerLR Sum_ICF v1.12 25OCT22</p> <p>REMAP-CAP_ProLR Sum_ICF v1.4 2022.10.25_tracked</p> <p>REMAP-CAP_ProLR Sum_ICF v1.4 25OCT22</p> <p>REMAP-CAP_UK_leaflet_V2.0_TC_2022.11.10</p> <p>REMAP-CAP_UK_leaflet_V2.0_2022.11.10</p> <p>REMAP-CAP_UK_poster_V2.0_TC_2022.11.09</p> <p>REMAP-CAP_UK_poster_V2.0_2022.11.09</p> <p>REMAP-CAP_YoungChild_PIS_ICF v1.0 25OCT22</p> <p>REMAP-CAP_Child_10-15_PIS_ICF v1.0 25OCT22</p> <p>REMAP-CAP_Child_16-17_PIS_ICF v1.0 25OCT22</p> <p>REMAP-CAP_UK_poster_Child V1.0_2022.11.09</p> <p>MhraProductsForm_AM037</p>	<ul style="list-style-type: none"> <li>• Submission of the Domain-Specific Appendix Antiviral domain (V2.0 tracked changes and final)- Addition of a new intervention, Baloxavir</li> <li>• Submission of the Domain-Specific Appendix corticosteroid domain (V5.0 tracked changes and final) - Addition of a new intervention, Dexamethasone</li> <li>• Changes to IS_CF for COVID-19, CAP and Influenza</li> <li>• Addition of IS_CF for children</li> <li>• Submission of adult and paediatric recruitment advertisement</li> <li>• Closure of the ACE2 RAS domain</li> <li>• Closure of the continuous therapeutic anticoagulation intervention in the prior anticoagulation domain</li> <li>• Pause of the Vitamin C domain</li> <li>• Extension to wards/ICU</li> </ul>
AM038	REC HRA MHRA	14.07.2023	10.08.2023	<p>REMAP-CAP AM038 Amendment_Tool_v1_6_LOCKED Contract_Issue_Letter_-_NIHR155209.docx (1)</p> <p>REMAP-CAP - Influenza Antiviral Domain-Specific Appendix - V2.1 - 14 Feb 2023_CLEAN</p> <p>REMAP-CAP - Influenza Antiviral Domain-Specific Appendix - V2.1 - 14 Feb 2023_TC from V2</p>	<ul style="list-style-type: none"> <li>• Change on DSA for antiviral '48hrs' not '96hrs'</li> <li>• Submission of a consent video</li> <li>• Submission of updated adult IS_CF</li> </ul>

				REMAP-CAP Text for ICU research video REMAP-CAP_PerLIS_ICF v1.14 2023.05.24_tracked REMAP-CAP_PerLIS_ICF v1.14 2023.05.24_clean REMAP-CAP_PerLR Sum_ICF v1.2 2023.05.24_tracked REMAP-CAP_PerLR Sum_ICF v1.2 2023.05.24_clean REMAP-CAP_PerLR Video Consent v1.1 2023.05.24_tracked REMAP-CAP_PerLR Video Consent v1.1 2023.05.24_clean REMAP-CAP_PIS_ICF v1.14 2023.05.24_tracked REMAP-CAP_PIS_ICF v1.14 2023.05.24_clean REMAP-CAP_ProLR Sum_ICF v1.6 2023.05.24_tracked REMAP-CAP_ProLR Sum_ICF v1.6 2023.05.24_clean REMAP-CAP_Video Consent v1.1 2023.05.24_tracked REMAP-CAP_Video Consent v1.1 2023.05.24_clean REMAP-CAP_SmPC_Amoxicillin-Clavulanate (Augmentin)_20200303 REMAP-CAP_SmPC_Azithromycin (Zedbac)_20180206 REMAP-CAP_SmPC_Azithromycin tablet_20220107 REMAP-CAP_SmPC_Ceftarolinefosamil (Zinforo)_20220215_stopped REMAP-CAP_SmPC_Ceftriaxone_20220119 REMAP-CAP_SmPC_Clarithromycin powder_20160914 REMAP-CAP_SmPC_Clarithromycin tablet_20171128 REMAP-CAP_SmPC_Erythromycin_20140425 REMAP-CAP_SmPC_Levofloxacin_20210106 REMAP-CAP_SmPC_Moxifloxacin (Avelox)_20201109 REMAP-CAP_SmPC_Piperacillin-Tazobactam (Tazocin)_20211223 REMAP-CAP_SmPC_Roxithromycin (RULID)_202005XX REMAP-CAP_SmPC_Hydrocortisone(Solu-Cortef)_20220107_eng REMAP-CAP_SmPC_DalteparinSodium(Fragmin)_20201009 REMAP-CAP_SmPC_Enoxaparin Sodium_20220427 REMAP-CAP_SmPC_Heparin Sodium_20180928 REMAP-CAP_SmPC_Tinzaparin Sodium_20210108 REMAP-CAP_SmPC_Dexamethasone Tablet_20220814_current REMAP-CAP_SmPC_Oseltamivir (Tamiflu)_20210525 REMAP-CAP_Privacy Notice v1.1 2023.04.18_tracked REMAP-CAP_Privacy Notice v1.1 2023.04.18_clean	<ul style="list-style-type: none"> <li>• Submission of the SmPC's for the antibiotic, macrolide, steroid, anticoagulation and antiviral domains</li> <li>• NIHR Grant agreement (extension of study end date)</li> </ul>
AM039	REC HRA MHRA	22.09.2023	21.11.2023	REMAP-CAP AM039 Amendment_Tool_v1_6_LOCKED Cysteamine_Nylexa IMPD_V5.1_27 Oct 2023_Final Cysteamine_Nylexa IMPD_V5.1_27 Oct 2023_track changed Cysteamine_Nylexa IMPD_Summary of changes_Oct 2023 REMAP-CAP_European RSA_Change History_20210910_eng_2 REMAP-CAP_Protocol_European RSA_V3.6_20201217_eng	<ul style="list-style-type: none"> <li>• IMPD update for cysteamine</li> <li>• European RSA V3.6</li> <li>• Addition of abx and macrolides to COVID pts in PIS</li> </ul>

				REMAP-CAP_Protocol_European_RSA_V3.6_20201217_eng_TC REMAP-CAP_PIS_ICF v1.15_19.09.2023 Clean REMAP-CAP_PIS_ICF v1.15_19.09.2023 TC REMAP-CAP_PerLIS_ICF v1.15_19.09.2023_Clean REMAP-CAP_PerLIS_ICF v1.15_19.09.2023_TC REMAP-CAP_PerLR Sum_ICF v1.3_19.09.2023_Clean REMAP-CAP_PerLR Sum_ICF v1.3_19.09.2023_TC REMAP-CAP_ProLR Sum_ICF v1.7_19.09.2023_Clean REMAP-CAP_ProLR Sum_ICF v1.7_19.09.2023_TC UK_REMAP-CAP_SubjectInsurance_Cert_20230110 DSMB_REMAP-CAP_20230808_ltr_No safety concerns	<ul style="list-style-type: none"> <li>Updated insurance certificate to Jun 2026</li> <li>DSMB letter</li> </ul>
AM040	22.12.2023	29.01.2024		REMAP-CAP AM040 Amendment_Tool_v1_6_LOCKED REMAP-CAP AM040 cover letter 22.12.2023 AM040_MhraProductsForm_snapshot REMAP-CAP letter to sites - Anticoagulation and Antiplatelet Domain - 30 November 2023 REMAP-CAP letter to sites - Corticosteroid Domain_06 December 2023 REMAP-CAP_Tel v2.0 2023.11.16_Clean REMAP-CAP_Tel v2.0 2023.11.16_Tracked REMAP-CAP_PerLR Sum_ICF v1.4_2023.11.16_Tracked REMAP-CAP_PerLR Sum_ICF v1.4_2023.11.16_Clean REMAP-CAP_PerLIS_ICF v1.16_2023.11.16_Tracked REMAP-CAP_PerLIS_ICF v1.16_2023.11.16_Clean REMAP-CAP_YoungChild_PIS_ICF v1.3 2023.12.13_Tracked REMAP-CAP_YoungChild_PIS_ICF v1.3 2023.12.13_Clean REMAP-CAP_ProLR Sum_ICF v1.8_2023.11.16_Tracked REMAP-CAP_ProLR Sum_ICF v1.8_2023.11.16_Clean REMAP-CAP_PIS_ICF v1.16_2023.11.16_Tracked REMAP-CAP_PIS_ICF v1.16_2023.11.16_Clean REMAP-CAP_Child_16-17_PIS_ICF v1.2 2023.11.16_Tracked REMAP-CAP_Child_16-17_PIS_ICF v1.2 2023.11.16_Clean REMAP-CAP_Child_10-15_PIS_ICF v1.4 2023.11.16_tracked REMAP-CAP_Child_10-15_PIS_ICF v1.4 2023.11.16_Clean REMAP-CAP pharmacy guide - Version 4 - 26 October 2023 REMAP-CAP Immune Modulation Influenza DSA - V1.0 Adult + Paediatric patients - 29 September 2023 REMAP-CAP Immune Modulation Influenza DSA - V1.1 Adult patients only - 29 September 2023	<ul style="list-style-type: none"> <li>Submission of the Immune Modulation DSA</li> <li>Updates to adult and children PIS documents (Telephone consent can now be used for legal representatives and patients)</li> <li>Adding postal consent option.</li> <li>Closure of fixed hydrocortisone intervention</li> <li>Closure of the anticoagulation and antiplatelet domains</li> <li>Submission of the pharmacy Manual v4.0</li> </ul>

AM41	REC HRA MHRA	20.05.2024	27.06.2024	<p>REMAP-CAP AM041 Amendment_Tool_v1_6_LOCKED</p> <p>REMAP-CAP letter to sites - Antibiotic and Macrolide Domains</p> <p>REMAP-CAP letter to sites - Cysteamine Domain</p> <p>Roche_REMAP-CAP_MA compliance Baloxavir</p> <p>UK_REMAP-CAP_SubjectInsurance_Cert_20240205</p> <p>UK_REMAP-CAP_SubjectInsurance_Endorsement 4_20240205</p> <p>Effective_UK (English) EQ-5D-Y Paper Interviewer Administration v1.3</p> <p>Effective_UK (English) EQ-5D-Y Paper Proxy2 Interviewer Administration v1.2</p> <p>REMAP-CAP-PerLIS-ICF-v1.18-07.05.2024 Clean</p> <p>REMAP-CAP-PerLIS-ICF-v1.18-07.05.2024-TC</p> <p>REMAP-CAP-PerLR-Sum-ICF-v1.6-07.05.2024- Clean</p> <p>REMAP-CAP-PerLR-Sum-ICF-v1.6-07.05.2024- TC</p> <p>REMAP-CAP-PIS-ICF-v1.18 07.05.2024 Clean</p> <p>REMAP-CAP-PIS-ICF-v1.18 07.05.2024 TC</p> <p>REMAP-CAP-ProLR-Sum-ICF-v1.10 07.05.2024 Clean</p> <p>REMAP-CAP-ProLR-Sum-ICF-v1.10 07.05.2024 TC</p> <p>GxP-0556690_MV43976_BALO_2FC_20MG_BL_39029775 Primary Packaging</p> <p>GxP-0556691_MV43976_BALO_2FC_20MG_BL_39029775_BFP</p> <p>Booklet front page for primary packaging</p> <p>GxP-0556692_MV43976_BALO_2FC_20MG_BX_39029774</p> <p>Secondary Packaging</p> <p>GxP-0556693_MV43976_BALO_2FC_20MG_BX_39029774_BFP</p> <p>Booklet frontpage for secondary packaging</p>	<ul style="list-style-type: none"> <li>• Change of baloxavir supply from Roche UK to Roche global</li> <li>• Addition of releasing site information to section D9 of the MHRA CTA for licensed Baloxavir</li> <li>• Submission of clinical trial labelling for baloxavir</li> <li>• Closure of the cysteamine domain</li> <li>• Closure of the antibiotics and macrolide domains for COVID patients.</li> <li>• Submission of the updated Insurance certificate</li> <li>• Matrons can act as ProLRs</li> <li>• Clarification that appropriately trained individuals can obtain consent</li> <li>• Submission of the EQ-5D-Y Questionnaire</li> </ul>
NS01	REC HRA			<p>Updated document: UK REMAPCAP PerLR ICF v2.3 2019.06.20</p> <p>Superseded document: UK REMAPCAP PerLR ICF v2.2 2019.06.20</p>	Notification of a typographical error in the REMAP-CAP information sheet / consent form for Personal Legal representative. On page 8, it was incorrectly titled ' consent form for patients able to give consent'. I have amended it to ' consent form for patients <b>unable</b> to give consent
NS02	REC HRA				Addition of 12 new recruiting sites (HRA and REC had confirmed that new sites should be added as part of non substantial amendment to fast track approvals during the pandemic)
NS03	REC			Old document	Typographical error on ProLR V1.1

	HRA MHRA			UK REMAP-CAP ProLR ICF V1.1 2018.11.30 Updated document UK REMAP-CAP ProLR ICF V1.2 2020.04.20	Header indicated Protocol version 2 rather than version 3
NS04	REC HRA	20.09.21	20.09.21	NIHR funding extension agreement	This involves clarification that the current funding allows for continued REMAP-CAP recruitment till the 30 <sup>th</sup> June 2022. Updated end date for patient recruitment. REMAP-CAP is designed as a perpetual study but additional funding extensions (provided by the RECOVER agreement and the NIHR) are now in place to continue recruitment of REMAP-CAP patients till 30.06.22
NS05	REC HRA	13.10.21	N/A	UK REMAP CAP Retro PIS ICF V2.6 2021.08.16 CLEAN	As part of AM028 all pandemic and non-pandemic participant information sheets and consent forms were updated, reviewed and approved, as well as the summary information sheets. Unfortunately due to the volume of participant information sheets submitted during the upload process the UK REMAP-CAP Retrospective Information for participants and Consent Form, V2.6 (track changed and clean version) dated 16th August 2021 was missed. The changes were exactly the same as those submitted, reviewed and approved for the other documentation submitted for English Sites, Northern Irish sites and Scottish sites. For this reason this missed document is being submitted as part of a non substantial amendment
NS06	HRA	07.08.2023	10.08.2023	REMAP-CAP_Child_10-15_PIS_ICF v1.3 2023.08.02	Changes to REMAP-CAP_Child_10-15_PIS_ICF



NS07	REC HRA	19.10.2023	27.10.2023	<p>Summary of changes to Corticosteroid Domain-Specific Appendix V5.0 - 16 October 2023</p> <p>Summary of changes to Influenza Antiviral Domain-Specific Appendix V2.2 - 16 October 2023</p> <p>REMAP-CAP Corticosteroid Domain-Specific Appendix V5.1_16 October 2023_CLEAN</p> <p>REMAP-CAP Corticosteroid Domain-Specific Appendix V5.1_16 October 2023_TC from V5</p> <p>REMAP-CAP - Influenza Antiviral Domain-Specific Appendix - V2.2 - 16 October 2023_CLEAN</p> <p>REMAP-CAP - Influenza Antiviral Domain-Specific Appendix - V2.2 - 16 October 2023_TC from V2.1</p> <p>DSMB_REMAP-CAP_20230922_itr_pediaticr incl and no safety concerns</p> <p>REMAP-CAP NSA07 Amendment Tool Locked</p>	<p>1. Update to the antiviral domain specific appendix - minor clarification to wording relating to paediatric participants</p> <p>2. Update to the corticosteroid domain specific appendix - minor clarification to wording relating to paediatric participants</p>
NSA08	HRA		05.12.2023	<p>REMAP-CAP_YoungChild_PIS_ICF v1.2 2023.11.13_Clean</p> <p>REMAP-CAP_YoungChild_PIS_ICF v1.2 2023.11.13_tracked</p>	<p>The parent information / consent was mistakenly removed whilst under review for AM037 – this has been reinstated.</p>
NSA09	HRA	16.04.2024	22.04.2024	<p>UK_REMAP-CAP_mNCA_V2.4_20207XX_Template</p> <p>UK_REMAP-CAP_mNCA_V2.4_20207XX_Template_TC</p> <p>UK REMAP-CAP poster_Child_V2.0 2024.03.18- TC</p> <p>UK REMAP-CAP poster_Child_V2.0 2024.03.18- CLEAN</p> <p>UK REMAP-CAP poster V2.1_2024.03.18 - TC</p> <p>UK REMAP-CAP poster V2.1_2024.03.18 - CLEAN</p> <p>REMAP-CAP-ProLR-Sum-ICF-v1.9-14.03.2024-CLEAN</p> <p>REMAP-CAP-ProLR-Sum-ICF-v1.9-14.03.2024-TC</p> <p>REMAP-CAP-PerLIS-ICF-v1.17-18.03.2024-CLEAN</p> <p>REMAP-CAP-PerLIS-ICF-v1.17-18.03.2024-TC</p> <p>REMAP-CAP-PerLR-Sum-ICF-v1.5-18.03.2024-TC</p> <p>REMAP-CAP-PerLR-Sum-ICF-v1.5-18.03.2024-CLEAN</p> <p>REMAP-CAP-PIS-ICF-v1.17-18.03.2024-TC</p> <p>REMAP-CAP-PIS-ICF-v1.17-18.03.2024-CLEAN</p> <p>REMAP-CAP-Child-10-15-PIS-ICF-v1.5-19.03.2024-TC</p> <p>REMAP-CAP-Child-10-15-PIS-ICF-v1.5-19.03.2024-CLEAN</p>	<ol style="list-style-type: none"> <li>1. New Contract (mNCA) template</li> <li>2. Updated adults and paediatric posters with QR code for new website.</li> <li>3. PIS document updates - PerLR and ProLR add space to add name of patient and also remove Farah's number from all PIS documents</li> <li>4. Acknowledgement of amendment still addressed to Ms. W. van Bentum-Puijk. Change this in Ms. S. Peters</li> <li>5. Clarification that consent forms can be sent by email or post</li> </ol>

Commented [FO1]: Check with Aisha

				REMAP-CAP-Child-16-17-PIS-ICF-v1.3-19.03.2024- TC REMAP-CAP-Child-16-17-PIS-ICF-v1.3-19.03.2024-CLEAN REMAP-CAP-Young-Child-PIS-ICF-v1.4-19.03.2024-TC REMAP-CAP-Young-Child-PIS-ICF-v1.4-19.03.2024-CLEAN	
NSA10	HRA	31.10.2024	31.10.2024	237150_NSA10_locked_PDF Grant agreement letter until Nov24	1. Correction of study end date to 30 <sup>th</sup> Nov 2024

**FYI, during the approval process for the 1<sup>st</sup> three substantial amendments, the HRA's incorrectly labelled every amendment as Amendment 01 on the approval letter. They were made aware of this, but as all the other pertinent information on the document was correct, they did not feel the need to re-issue the letter.**