

Supplementary appendix

This appendix formed part of the original submission and has been peer reviewed. We post it as supplied by the authors.

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Supplementary appendix

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Supplementary Table S1 Ongoing registered trials of immunomodulation therapy for COVID-19

| Strategy | Intervention | Trial registration # | Sponsor | Registration date | Study design |
|-----------------|----------------------------------------------------------------------------------------------------|-------------------------|------------------------------------------------------------------------------------------------------------|-------------------|----------------------------------------------------------------|
| IL-6 antagonism | Tocilizumab | ChiCTR2000029765 | The First Affiliated Hospital of University of science and technology of China (Anhui Provincial Hospital) | 2020-02-13 | multicenter, randomized controlled trial |
| | Favipiravir + Tocilizumab Favipiravir Tocilizumab | ChiCTR2000030894 | Peking University First Hospital | 2020-03-16 | Multicenter, Randomized, Controlled |
| | tozumab and adamumab | <u>ChiCTR2000030580</u> | Shanghai General Hospital | 2020-03-08 | randomized controlled |
| | Tocilizumab: 8 mg/kg injection | NCT04317092 | National Cancer Institute, Naples | 2020-03-20 | open-label, single- group |
| | Tocilizumab: 400mg | NCT04310228 | Peking University First Hospital | 2020-03-17 | open-label, randomized |
| | Tocilizumab | NCT04332094 | Fundació Institut de Recerca de l'Hospital de la Santa Creu i Sant Pau | 2020-04-02 | Randomized, Multicenter, Open- label |
| | Anakinra: 100 mg/d, sc, 28 d Siltuximab: 11 mg/kg, single IV Tocilizumab: 100 mg/d, sc, 28 d | NCT04330638 | University Hospital, Ghent | 2020-04-01 | randomized, open- label |
| | Tocilizumab: 8 mg/kg | NCT04315480 | Università Politecnica delle Marche | 2020-03-19 | Open-label, single arm |
| | Tocilizumab: 200mg, single dose Tocilizumab: 80mg, single dose | NCT04331795 | University of Chicago | 2020-04-02 | open-label, non- randomized |
| | Tocilizumab 8mg/kg | NCT04331808 | Assistance Publique - Hôpitaux de Paris | 2020-04-02 | randomized, open- label, controlled |
| | Tocilizumab: 8 mg/kg, one dose | NCT04320615 | Hoffmann-La Roche | 2020-03-25 | randomized, multicenter, double- blind, controlled trial |
| | Sarilumab: low dose Sarilumab high dose | NCT04315298 | Regeneron Pharmaceuticals | 2020-03-19 | Randomized, Double-Blind, Controlled |
| IL-1 blockade | Anakinra Emapalumab | NCT04324021 | Swedish Orphan Biovitrum | 2020-03-27 | multicenter, randomized, open- label |
| | Anakinra: 100 mg/d, sc, 28 d | NCT04330638 | University Hospital, Ghent | 2020-04-01 | randomized, open- |

| | Siltuximab: 11 mg/kg, single IV Tocilizumab: 100 mg/d, sc, 28 d | | | | label |
|-----------------------------------|------------------------------------------------------------------------------------------------------------------------------------|------------------|----------------------------------------------------------------------------------------------|------------|----------------------------------------------------------------|
| Pyrimidine synthesis inhibitor | leflunomide | ChiCTR2000030058 | Renmin Hospital of Wuhan University | 2020-02-22 | multicenter, randomized, double- blind, controlled trial |
| JAK inhibitor | Ruxolitinib combined with mesenchymal stem cell | ChiCTR2000029580 | Tongji Hospital, Tongji Medical College, Huazhong University of Science and Technology | 2020-02-05 | prospective, single- blind, randomized controlled trial |
| | Ruxolitinib 10 mg twice a day | NCT04334044 | Grupo Cooperativo de Hemopatías Malignas | 2020-04-03 | Open-label, single group |
| | Ruxolitinib | NCT04331665 | University Health Network, Toronto | 2020-04-02 | Single Arm Open- label |
| | Baricitinib 4 mg/day x 14 days | NCT04320277 | Hospital of Prato | 2020-03-24 | open label, non- randomized |
| | Tofacitinib 10mg bid x 14 days | NCT04332042 | Università Politecnica delle Marche | 2020-04-02 | Single Arm Open- label |
| Thalidomide | Thalidomide: 100mg, po, qn, for 14 days | NCT04273529 | First Affiliated Hospital of Wenzhou Medical University | 2020-02-20 | Randomized |
| | Thalidomide: 100mg, po, qn, for 14 days | NCT04273581 | First Affiliated Hospital of Wenzhou Medical University | 2020-02-18 | Randomized |
| Cloroquine/ Hydroxychloroquine | Chloroquine phosphate, Hydroxychloroquine sulfate | ChiCTR2000030054 | Zhongshan Hospital Affiliated to Xiamen University | 2020-02-22 | prospective, open label, randomized, controlled trial |
| | hydroxychloroquine | NCT04333225 | Baylor Research Institute | 2020-04-03 | open label, non- randomized |
| | Hydroxychloroquine 200mg, q12h, 7-10 days | NCT04307693 | Asan Medical Center | 2020-03-13 | Multicenter, open labelled, randomized |
| | Hydroxychloroquine: 400mg/day for 5 day | NCT04261517 | Shanghai Public Health Clinical Center | 2020-02-07 | randomized, open- label |
| | Hydroxychloroquine | NCT04328467 | University of Minnesota | 2020-03-31 | randomized |
| | hydroxychloroquine 400 mg bid for 1 day, 200 mg bid for 4 days azithromycin 500 mg on day 1 plus 250 mg daily on days 2-5 | NCT04329832 | Intermountain Health Care, Inc. | 2020-04-01 | randomized, open- label |
| | hydroxychloroquine 400 mg bid for 1 day, 200 mg bid for 4 days | NCT04329611 | University of Calgary | 2020-04-01 | Randomized, Double- blind, controlled |
| | Hydroxychloroquine | NCT04325893 | University Hospital, Angers | 2020-03-30 | Randomized, Double-blind |
| | hydroxychloroquine 400 mg bid | NCT04329572 | Azidus Brasil | 2020-04-01 | open-label, single |

| for 1 day, 200 mg bid for 4 days | | | | group | |
|-------------------------------------------------------------------------------------------------------------------------|-------------|----------------------------------------------------------------------------|------------|--------------------------------------------------------------------|--|
| hydroxychloroquine 400 mg bid for 1 day, 200 mg bid for 9 days | NCT04323631 | Rambam Health Care Campus | 2020-03-26 | randomized, open- label | |
| Hydroxychloroquine + azithromycin Hydroxychloroquine | NCT04321278 | Hospital Israelita Albert Einstein | 2020-03-25 | randomized, open- label | |
| hydroxychloroquine 400mg day for 10 days | NCT04315896 | National Institute of Respiratory Diseases, Mexico | 2020-03-20 | Double blinded, randomized controlled trial | |
| hydroxychloroquine: 300mg weekly hydroxychloroquine: 300mg twice weekly hydroxychloroquine: 150 mg | NCT04333732 | Washington University School of Medicine | 2020-04-03 | multi-site, randomized, double- blinded, controlled trial | |
| daily hydroxychloroquine or chloroquine: 10 mg / kg followed by 155 mg daily | NCT04303507 | University of Oxford | 2020-03-11 | randomised, controlled | |
| hydroxychloroquine 400 mg bid for 1 day, 200 mg bid for 4 days Azithromycin 500 mg for 1day, 250 mg for 4 days | NCT04329832 | Intermountain Health Care, Inc. | 2020-04-01 | randomized, open- label | |
| Hydroxychloroquine | NCT04328272 | Prof. Dr. Umar Farooq | 2020-03-31 | Single blinded randomized | |
| Hydrochloroquine 400 mg x 3 days, then 200 mg x 11 days | NCT04328961 | University of Washington | 2020-04-01 | Single blinded randomized | |
| Chloroquine | NCT04328493 | Oxford University Clinical Research Unit, Vietnam | 2020-03-31 | open label, randomized, controlled trial | |
| hydroxychloroquine 400 mg bid for 1 day, 200 mg bid for 4 days | NCT04332991 | Massachusetts General Hospital | 2020-04-03 | Randomized | |
| chloroquine 500 mg bid for 7 days | NCT04333628 | HaEmek Medical Center, Israel | 2020-04-03 | open label, | |
| | | University of Minnesota | 2020-03-16 | Randomized | |
| Hydroxychloroquine | NCT04333654 | Sanofi | 2020-04-03 | Randomized, double- blinded, controlled | |
| Hydroxychloroquine | NCT04322396 | Chronic Obstructive Pulmonary Disease Trial 2020-03-26 Network, Denmark | | Randomized, double- blinded, controlled | |

| Chloroquine 1.0gx2 days, 0.5gx12 day from the third day, Hydroxychloroquine 0.2g bid x 14 days | ChiCTR2000029992 | Zhongshan Hospital Affiliated to Xiamen University | 2020-02-18 | prospective, randomized, open label, controlled trial |
|------------------------------------------------------------------------------------------------|------------------|--------------------------------------------------------------------------------------------------|------------|-------------------------------------------------------------|
| Hydroxychloroquine: Day1-3: 500mg BID, Day4-10: 250mg BID | ChiCTR2000029899 | Peking University Third Hospital | 2020-02-16 | Randomized, Open- label, Parallel, Controlled Trial |
| Hydroxychloroquine: Day1-3: 500mg BID, Day4-10: 250mg BID | ChiCTR2000029898 | Peking University Third Hospital | 2020-02-16 | Randomized, Open- label, Parallel, Controlled Trial |
| Hydroxychloroquine | ChiCTR2000029868 | Ruijin Hospital, Shanghai Jiaotong University School of Medicine | 2020-02-15 | multicenter, randomized controlled trial |
| Hydroxychloroquine | ChiCTR2000029803 | Renmin Hospital of Wuhan University | 2020-02-14 | prospective, randomized, open- label, controlled |
| Hydroxychloroquine | ChiCTR2000029762 | The First Affiliated Hospital of Chongqing Medical University | 2020-02-12 | randomized, open- label, controlled trial |
| Hydroxychloroquine: low-, medium-, high-dose | ChiCTR2000029761 | The First Affiliated Hospital of Chongqing Medical University | 2020-02-12 | Multicenter, randomized, open- label |
| Hydroxychloroquin | ChiCTR2000029760 | The Second Affiliated Hospital of Chongqing Medical University | 2020-02-12 | randomized controlled study |
| Hydroxycholoroquine: 0.2g twice a day | ChiCTR2000029740 | The First Hospital of Peking University | 2020-02-11 | randomized open- label controlled |
| Hydroxychloroquine: 0.2g oral 2/ day Hydroxychloroquine 0.1g oral 2/ day | ChiCTR2000029559 | Renmin Hospital of Wuhan University | 2020-02-04 | Randomized, double blind, Parallel |
| phosphoric chloroquine: 2 tablets BID | ChiCTR2000030031 | The Sixth Affiliated Hospital of Guangzhou Medical University (Qingyuan People's Hospital) | 2020-02-21 | randomized, double- blind, parallel, controlled trial |
| phosphoric chloroquine: 2 tablets BID | ChiCTR2000029837 | Jingzhou Central Hospital | 2020-02-15 | randomized, double- blind, parallel, controlled trial |
| chloroquine | ChiCTR2000031204 | Beijing you'an Hospital, Capital Medical University | 2020-03-24 | randomized |
| chloroquine | ChiCTR2000030718 | Zhongnan Hospital of Wuhan University | 2020-03-11 | open label randomized |
| favipiravir plus chloroquine | ChiCTR2000030987 | Beijing Chao-yang Hospital, Capital Medical | 2020-03-20 | randomized |

| | favipiravir | | University | | controlled |
|-------------------|--------------------------------------------------------------------|-------------------------|----------------------------------------------------------------------------------------------|------------|----------------------------------------------------------------------|
| | phosphoric chloroquine: 2 tablets BID | ChiCTR2000029826 | Jingzhou Central Hospital | 2020-02-14 | randomized, double- blind, parallel, controlled trial |
| | Chloroquine Phosphate | ChiCTR2000029988 | Zhongnan Hospital of Wuhan University | 2020-02-18 | |
| | chloroquine phosphate aerosol inhalation | ChiCTR2000029975 | The First Hospital of Jilin University | 2020-02-18 | Open label, Parallel Single arm |
| | Chloroquine Phosphate | ChiCTR2000029939 | HwaMei Hospital, University of Chinese Academy of Sciences | 2020-02-16 | Single-blind, Randomized, Controlled |
| | Chloroquine Phosphate | ChiCTR2000029935 | HwaMei Hospital, University of Chinese Academy of Sciences | 2020-02-16 | Single arm |
| | Chloroquine Phosphate vs. Lopinavir / Ritonavir | ChiCTR2000029741 | The Fifth Affiliated Hospital Sun Yat-Sen University | 2020-02-11 | prospective, open- label, multicenter randomized controlled |
| | chloroquine | ChiCTR2000029609 | The Fifth Affiliated Hospital of Sun Yat-Sen University | 2020-02-06 | prospective, open- label, multiple-center study |
| | chloroquine | ChiCTR2000029542 | Sun Yat sen Memorial Hospital of Sun Yat sen University | 2020-02-03 | prospective |
| Stem cell therapy | Iv injection of Wharton's Jelly mesenchymal stem cells (1×10^6/kg) | ChiCTR2000030088 | The Sixth Medical Center of PLA General Hospital | 2020-02-22 | Parallel |
| | Mesenchymal stem cell therapy | ChiCTR2000030020 | Second Hospital of University of South China, Hengyang | 2020-02-20 | Sequential |
| | mesenchymal stem cell therapy | ChiCTR2000029990 | Institute of Basic Medical Sciences, Chinese Academy of Medical Sciences | 2020-02-18 | Parallel |
| | Human Menstrual Blood- Derived Stem Cells | <u>ChiCTR2000029606</u> | The First Affiliated Hospital, College of Medicine, Zhejiang University | 2020-02-07 | open label, Parallel |
| | Ruxolitinib combined with mesenchymal stem cell | ChiCTR2000029580 | Tongji Hospital, Tongji Medical College, Huazhong University of Science and Technology | 2020-02-05 | prospective, single- blind, randomized controlled trial |
| | Cord Blood Mesenchymal Stem Cell | ChiCTR2000029816 | Guangzhou reborn health management consultation co., LTD | 2020-02-14 | open label, Parallel |
| | umbilical cord mesenchymal stem cell conditioned medium | ChiCTR2000029569 | Xiangyang First People's Hospital | 2020-02-04 | randomized controlled |

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|-------------------------------|--------------------------------------------------------|------------------|-----------------------------------------------------------------------------------------------|------------|-------------------------------------------------------|
| | mesenchymal stem cells | ChiCTR2000030835 | the First Affiliated Hospital of Xinxiang Medical University | 2020-03-15 | Non-randomized |
| | mesenchymal stem cells | ChiCTR2000031430 | The Fifth Medical Center of PLA General Hospital | 2020-03-31 | non-randomized controlled |
| | Dental pulp mesenchymal stem cells | ChiCTR2000031319 | Renmin Hospital of Wuhan University | 2020-03-27 | randomized controlled |
| | mesenchymal stem cells | ChiCTR2000031494 | Huangshi Hospital of Traditional Chinese Medicine | 2020-04-02 | randomized controlled |
| | mesenchymal stem cell | NCT04252118 | Beijing 302 Hospital, China | 2020-02-05 | open label, non- randomized |
| | Cord Blood Mesenchymal Stem Cell | NCT04269525 | ZhiYong Peng, Zhongnan Hospital | 2020-02-13 | Open-label, single arm |
| | Umbilical cord mesenchymal stem cells | NCT04273646 | Wuhan Union Hospital, China | 2020-02-18 | open label, randomized |
| Convalescent plasma treatment | Convalescent Plasma | NCT04333355 | Hospital San Jose Tec de Monterrey | 2020-04-03 | Open-label, single group |
| 1 | Convalescent Plasma | NCT04327349 | Mazandaran University of Medical Sciences | 2020-03-31 | Open-label, single arm |
| | Immunoglobulin of cured patients | ChiCTR2000030841 | Union hospital of Tongji Medical College, Huazhong University of Science and Technology | 2020-03-16 | randomized control |
| | Convalescent Plasma | ChiCTR2000030627 | The First Affiliated Hospital of Zhengzhou University | 2020-03-08 | randomized controlled |
| | infusing convalescent plasma: 200-500ml, two infusions | ChiCTR2000030039 | Affiliated Hospital of Xuzhou Medical University | 2020-02-13 | Multi-center, non randomized control |
| | anti-2019-nCoV inactivated convalescent plasma | ChiCTR2000030046 | First People's Hospital of Jiangxia District, Wuhan (Union Jiangnan Hospital) | 2020-02-21 | Multi-center, single arm trial |
| | anti-SARS-CoV-2 virus inactivated plasma | ChiCTR2000030010 | Wuhan Jinyintan Hospital (Wuhan Infectious Diseases Hospital) | 2020-02-19 | randomized, double- blind, parallel- controlled |
| | convalescent plasma treatment | ChiCTR2000029850 | The First Affiliated Hospital of Zhejiang University School of Medicine | 2020-02-15 | Single center, Non randomized control |
| | Convalescent plasma | ChiCTR2000029757 | Institute of Blood Transfusion, Chinese Academy of Medical Sciences | 2020-02-12 | prospective randomized controlled trial |
| Glucocorticoid | Methylprednisolone 250mg/24h | NCT04329650 | Judit Pich Martínez | 2020-04-01 | Randomized, open- label |
| | Dexamethasone | NCT04325061 | Dr. Negrin University Hospital | 2020-03-27 | randomized, controlled, open-label trial |

| Methylprednisolone: 80 mg/d | NCT04323592 | University of Trieste | 2020-03-26 | single arm, non |
|-------------------------------------------------|------------------|------------------------------------------------|------------|----------------------------|
| dexamethasone 20mg/d x 5 days + 10mg/d x 5 days | NCT04327401 | Luiz F. L. Reis, Ph.D., Hospital Sirio-Libanes | 2020-03-31 | randomized, open- label |
| Early corticosteroid Middle-late corticosteroid | ChiCTR2000030481 | Zhongnan Hospital of Wuhan University | 2020-03-03 | randomized control |
| low-dose methylprednisolone | ChiCTR2000029656 | Wuhan Pulmonary Hosptial | 2020-02-09 | randomized, open- label |

Trials listed are from ChiCtr.org.cn/ and ClinicalTrials.gov/, as of 2020-04-03.