Comprehensive overview of COVID-19 clinical trials
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Abstract
Here, we present an overview of the clinical trials that are currently being conducted or have concluded to date on COVID-19 globally. A comprehensive search was conducted to present 16 trial registries from around the world. Collectively, there are 1,528 trials reported for COVID-19 to date. Out of them, 50 studies included paediatric age group from day 0 to less than or equal to 18 years of age. A few 18 studies involve only females and 20 only males. There are 2 trials currently underway in Bangladesh, 4 in Pakistan and 13 in India. Overall, 940 trials are related to medicines and/or interventions. They include standard of care for any viral illness, antivirals, anti-inflammatory and immune altering medications. Two out of 10 vaccine trials are novel vaccines. It is knowledgeable and resourceful to reach out to the concerned sponsor if a physician thinks his patient can benefit from the trials in the region.

Keywords: Coronavirus, 2019-nCoV, SARS-CoV-2, 2019nCoV, Clinical Trial, Asia.

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Introduction
There are numerous speculations about diagnosis and treatment of the novel coronavirus detected in the end of 2019 (COVID-19) since Wuhan’s whistleblowers informed the world about the current outbreak.1 As members of the scientific community we need to inform our patients and communities with sound scientific information. Hence, we present here an overview of the clinical trials that are currently being conducted or have concluded to date on COVID-19 globally.

Methods
To have an extensive search, 16 trial registries from around the world were included. These include Australian New Zealand Clinical Trials Registry (ANZCTR),2 Chinese Clinical Trial Registry (ChiCTR),3 ClinicalTrials.gov from United States,4 Clinical Trials Registry - India (CTRI),5 European Union (EU) Clinical Trials Register,6 DRKS - German Clinical Trials Register,7 Iranian Registry of Clinical Trials,8 International Standard Randomized Controlled Trial Number (ISRCTN)9 registry by World Health Organization (WHO), International Clinical Trials Registry Platform (ICTRP) and the International Committee of Medical Journal Editors (ICMJE), Japan Primary Registers Network (JPRN),10 Lebanese Clinical Trials Registry (LBCTR),11 Netherlands Trial Registry (NTR),12 Pan African Clinical Trial Registry (PACTR),13 Brazilian Registry of Clinical Trials (ReBEC),14 Peruvian Clinical Trials Registry (REPEC),15 Cuban Public Registry of Clinical Trials (RPCEC),16 and Thai Clinical Trials Registry (TCTR).17 All these registries combine their data under the umbrella of WHO to ensure a single point of access and the documentation of trials.18 We used it to extract all our findings. The data is dynamic and updated frequently. Hence, we report all findings as noted on April 27, 2020.18

Results
Demographics
Collectively there are 1,528 trials involving COVID-19 to date.18 Fifty studies included paediatric age group from day 0 to less than or equal to 18 years of age. Two studies involved newborns up to 1 year of age currently underway in China, of which one is an interventional study testing moxibustion in preventing COVID-19 pneumonia. The other is an observational study about pneumonia due to COVID-19 in neonates.18

There are 18 studies involving only females.18 Majority (n=14) are just observational and are related to pregnancy. A researcher from China is studying the impact of the novel virus on ovarian function as a basic science project.18 A study about diagnostic testing such as detecting the virus in vaginal secretions is also currently underway in China.18 Two studies are interventional: one related to postpartum depression and another about effects of a mobile meditation application on stress. There are 20 studies involving only males.18 Most (n=14)
### Table 1: Details of trials planned for some Asian countries

<table>
<thead>
<tr>
<th>Countries</th>
<th>Source Register</th>
<th>Scientific Title</th>
<th>Primary Sponsor</th>
<th>Study Type</th>
<th>Phase</th>
</tr>
</thead>
<tbody>
<tr>
<td>Afghanistan; Bangladesh; Bangladesh; Bhutan; India; Maldives; Pakistan; Sri Lanka</td>
<td>ISRCTN</td>
<td>The Impact of The COVID-19 Pandemic On The Provision, Practice, And Outcomes Of Vascular Surgery. An International Cohort Study (COVER)</td>
<td>University Hospitals Coventry and Warwickshire NHS Trust</td>
<td>Observational</td>
<td>N/A</td>
</tr>
<tr>
<td>Bangladesh</td>
<td>CTRI</td>
<td>Knowledge, Attitudes, And Fear Of COVID-19 During The Rapid Rise Period In Bangladesh</td>
<td>K M Amran Hossain</td>
<td>Observational</td>
<td>N/A</td>
</tr>
<tr>
<td>India</td>
<td>REPEC</td>
<td>Solidarity: An International Randomized Controlled Trial To Evaluate Non-Licensed Covid-19</td>
<td>OMS/OPS, Coventry and Warwickshire NHS Trust</td>
<td>Interventional</td>
<td>N/A</td>
</tr>
<tr>
<td>India</td>
<td>Clinicaltrials.Gov</td>
<td>A Clinical Trial To Evaluate The Safety And Efficacy Of Mycobacterium W In Critically Ill Patients Suffering From COVID 19 Infection</td>
<td>Cadila Pharmaceuticals</td>
<td>Interventional</td>
<td>N/A</td>
</tr>
<tr>
<td>India</td>
<td>CTRI</td>
<td>Max COVID Study</td>
<td>Max Super Specialty Hospital A Unit Of Devki Devi Foundation</td>
<td>Observational</td>
<td>N/A</td>
</tr>
<tr>
<td>India</td>
<td>CTRI</td>
<td>Open Labelled Randomised Controlled Trial To Study The Effect Of Chloroquine In Addition To Standard Therapy In COVID-19 Patients</td>
<td>Command Hospital Airforce</td>
<td>Interventional</td>
<td>N/A</td>
</tr>
<tr>
<td>India</td>
<td>CTRI</td>
<td>Outbreak Of Respiratory Tract Infection Score (ORTIS); Objective Screening For Children To Rule Out COVID-19 And Prevent Nosocomial Spread - ORTIS</td>
<td>DACH Jaipur</td>
<td>Observational</td>
<td>N/A</td>
</tr>
<tr>
<td>India</td>
<td>CTRI</td>
<td>Viral Infection And Respiratory Illness Universal Study: COVID-19 Registry And Validation Of C2D2 (Critical Care Data Dictionary)</td>
<td>Discovery The Critical Care Research Network</td>
<td>Observational</td>
<td>N/A</td>
</tr>
<tr>
<td>India</td>
<td>Clinicaltrials.Gov</td>
<td>Efficacy Of Convalescent Plasma Therapy In Severely Sick COVID-19 Patients: A Pilot Randomized Controlled Trial</td>
<td>Institute Of Liver And Biliary Sciences, India</td>
<td>Interventional</td>
<td>Phase 2</td>
</tr>
<tr>
<td>India</td>
<td>EU Clinical Trials Register</td>
<td>Preventing Pulmonary Complications In Surgical Patients At Risk Of COVID-19 - Covidsurget_V1.0_20200330</td>
<td>University Of Birmingham</td>
<td>Interventional</td>
<td>Phase 3</td>
</tr>
<tr>
<td>India</td>
<td>CTRI</td>
<td>Hydroxychloroquine, In Open Labelled, Randomised Intervention For Prevention Of New Infection And Adverse Outcomes Following COVID-19 Infection- A Tertiary Hospital Based Study - CORONA Study</td>
<td>Dr Remesh Bhasi</td>
<td>Interventional</td>
<td>Phase 3</td>
</tr>
<tr>
<td>India</td>
<td>Clinicaltrials.Gov</td>
<td>An Open Labelled Trial To Evaluate Safety And Efficacy Of SSV Formulation To Boost Immunity In Quarantine Patients Of COVID-19</td>
<td>SSV Phytopharmaceuticals</td>
<td>Interventional</td>
<td>Phase 3</td>
</tr>
<tr>
<td>India</td>
<td>Clinicaltrials.Gov</td>
<td>A Prospective, Longitudinal, Observational Study Of Healthcare Workers And The General Population To Watch For Flu-Like Symptoms In Suspicion Of COVID 19</td>
<td>Max Healthcare Institute Limited</td>
<td>Observational</td>
<td>N/A</td>
</tr>
</tbody>
</table>

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of which are observational while only one is a single validation study about COVID-19 RNA diagnostic kits among males happening in China. Remaining trials included both genders.

There are 2 trials in Bangladesh, 4 in Pakistan and 13 in India. Details of trials currently underway in some Asian countries are mentioned in Table-1.

**Treatment Trials**

Scientists are studying treatment options from a vast array of new medications to household remedies noted in various cultures. Overall, 940 trials are related to medicines and/or interventions. The top treatment options currently being studied are considerable.\(^1\),\(^2\),\(^3\) They include standard care for any viral illness, antivirals, anti-inflammatory and immune altering medications. Direct-acting antiviral drugs, LPV/r (lopinivir-ritonavir), LPV/r plus direct-acting antiviral drugs, LPV/r plus interferon, LPV/r plus corticosteroids, favipiravir, remdesivir, umifenovir, umifenovir plus interferon, azithromycin, tocilizumab are the most tested immunosuppressive drugs.\(^1\),\(^2\),\(^3\) Others include sarilumab, corticosteroids and hydroxychloroquine plus antivirals. A total of 157 trials include chloroquine phosphate and/or hydroxychloroquine sulphate,\(^1\),\(^2\) used either alone or in combination with another medication. ClinicalTrials.gov reported that a Brazilian study was terminated at early stages due to high death rates associated with the adverse effects linked to higher doses of hydroxychloroquine and azithromycin required to treat the novel virus.\(^2\),\(^3\)

Seven immunoglobulin and 41 inactivated convalescent plasma and some stem cell therapies are also being tested.\(^1\) Psychological interventions are also being studied for adjacent therapy.

**Vaccine Trials**

To date, 10 vaccine trials are underway.\(^1\) Two are novel LV-SMENP-DC and Cuban vaccine CIGB 2020 (CIGB 2020).\(^1\) Others are previously used vaccines that are being tested for their safety and efficacy on COVID-19 prevention. These include Bacille Calmette-Guérin (BCG), adenovirus type 5, measles, mumps, and rubella (MMR).

Traditional Chinese Medicine is the most studied other than allopathic medicines. Twenty-eight are traditional, mostly Chinese such as Lianhua Qingwen Capsules.\(^1\) Others are from Iran including medium-chain triglyceride intake and a mixture of Astragalus gossypinus plus food spice based on Ferula assa-foetida mixed in apple juice.

**Conclusion**

This is an overview of the clinical trials underway to face this pandemic. It is knowledgeable and resourceful to reach out to the concerned sponsor if a physician thinks that his patient might benefit from the trials in the region.

**References**

2. Australian New Zealand Clinical Trials Registry [Internet] Sydney


