

Version	Date	SIGNIFICANT REVISIONS (compared to previous)
V11	April 20, 2022	<ul style="list-style-type: none"> • Outpatient management: reference to “pre-emptive therapy” instead of "for monoclonal antibody or antiviral" • Decision tree for pre-emptive therapy: sotrovimab no longer recommended (not effective at neutralizing the BA2 variant of SARS-CoV2 (currently predominant strain in Quebec). Main offerings are now Paxlovid (po) or remdesivir IV x 3d. • Paxlovid to be prescribed by the treating (MRP) physician (ID-HOT not needed if there are no contraindications for this drug). Links to a drug interaction checker and to the MSSS template prescription (contains the required RAMQ codes) are provided • Remdesivir: ID-HOT needed to give first dose in Medical day (barring staffing issues), and doses 2 and 3 via CLSC • Inpatient management of COVID: ID no longer needs to approve Remdesivir prescriptions in mild or moderate disease • For severe disease: sarilumab replaced with baricitinib, because of paucity of data showing mortality benefit for sarilumab - baricitinib now the preferred alternative to tocilizumab if this product is unavailable. ID consult still required for severe disease • Footnotes updated.
V10	Jan 23, 2022	<ul style="list-style-type: none"> • Outpatient management: hyperlink to a decision tree for selection of candidates for Monoclonal antibody (Sotrovimab) vs oral antiviral (paxlovid) • Addition of paxlovid as an option (with pharmacy consultation) • Option of budesonide for outpatients not meeting criteria for MAb/paxlovid.

V9	Dec 21, 2021	<ul style="list-style-type: none"> • Remdesivir: recent evidence suggest it may have more benefit than initially thought (time to recovery and progression to severe), even in the group who received it after a longer/unclear duration of symptoms. Remdesivir recommended for <u>all moderate</u> disease (rather than only if <7d since onset) • Monoclonal antibodies: specifying that Sotrivimab should be used for omicron, and Casrivimab/imdevimab for delta - ID still needs to be involved in all cases. • For receipt of Monoclonal antibodies, have defined the group "predicted to have sub-optimal response to vaccination" as: immunocompromised state with a list of conditions (including hypogammaglobulinemia) • Tocilizumab: global shortage – need to save on vials. Now recommending to switch to a fixed dose (400mg if <75kg; 600mg if 75-99kg; 800mg if > 100kg (rather than 8mg/kg); also suggesting Sarilumab single dose (another IL-6 inhibitor) as an alternative if we run out of tocilizumab. • Cytokine storm criteria removed: this was initially to guide use of Tocilizumab but proven too complex and was not used in any of the studies that showed a benefit with IL-6 inhibitors. Suggesting to focus on ruling out bacterial infection and only using CRP cut-off in cases that are already severe (by resp criteria) • baricitinib an option (but very limited stock) if no IL-6 inhibitors unavailable • Anticoagulation: LMWH dosage recommendations for obesity, pregnancy and eGFR 20-30; for eGFR <20 caution advised; UFH remains nursing intensive and presents Infection control issues – unclear risk/benefit in that population.
V8	Oct 13, 2021	<ul style="list-style-type: none"> • "Suggested admission criteria": removed the criterion stating “predicted to progress to severe disease if managed as an outpatient” since now offering Monoclonal antibodies (either through ID Day-Hospital or in-patient) • Management of mild disease now divided into 2 categories: low risk of progression to severe vs high risk of progression to severe. If high risk, AND presenting within <7d of symptoms --> candidates for Monoclonals (with ID approval); • Management of Moderate disease: MAb to be considered in those within <7d of symptoms and not vaccinated or reason to believe suboptimal response to vaccination • Footnotes related to MAb product.
V7	Feb 9, 2021	<ul style="list-style-type: none"> • We now recommend to <u>therapeutically anticoagulate mild-moderate cases</u> (rather than prophylactic anticoagulation) – this is based on recent data from 3 international trials, which showed a marked benefit (reduced severity of illness) with therapeutic anticoagulation. <u>No benefit and possible harm noted in severe cases.</u> Though the studies are not published, the guideline working group reached consensus after reviewing data provided by the PIs of the studies. <u>The list of contra-indications to therapeutic anticoagulation is deliberately conservative ie. more cautious than when treating confirmed DVT/PE.</u> • Some clarification around Tocilizumab use (and dosage provided)
V6	Jan 5, 2021	<ul style="list-style-type: none"> • <u>Admission criteria:</u> among the non-respiratory criteria, replaced our previous “Hematopoietic transplant recipients (HSCT) <u>with</u> high Immunodeficiency Scoring Index ISI*, or HIV with CD4 < 200” with a more generic one stating: “Concern for high risk of complications/severe disease if managed in outpatient - based on comorbidities* and living situation”, and added a box listing the comorbidities known to be associated with severe disease (in accordance with CDC and PHAC guidelines). • <u>Empiric Antibiotics:</u> <ul style="list-style-type: none"> • NO empiric antibiotics for moderate disease. There is now sufficient evidence that rates of superinfection are very low in COVID-19, and there has been significant overprescribing of antibiotic.

		<ul style="list-style-type: none"> • For patients with severe disease (high flow or mechanical ventilation), empiric antibiotics acceptable: suggest to cover community pathogens with ceftriaxone if hospitalized < 5-days, and broaden coverage with pip-tazo if hospitalized > 5-days • no need for azithro or doxy given data showing no benefit of coverage for atypicals in COVID-19 • <u>New criteria for prediction of COVID-cytokine storm</u> added under “additional considerations”: contrary to other criteria/risk calculators (H-Score, MAS score etc), these correlate well with clinical consensus of severe hyperimmune response; based on a recent publication (https://ard.bmj.com/content/80/1/88).
V5	Sept 30, 2020	<ul style="list-style-type: none"> • Provided criteria for use of Remdesivir outside of clinical trials (only for moderate disease AND < 10 days of symptoms – need ID approval; close monitoring of renal and hepatic functions) • Clarified classification as mild (O₂ sat > 92% on RA), moderate (need supplemental O₂ for O₂Sat > 92%) and severe disease (need high-flow or mechanical ventilation to maintain O₂ sat > 92%) • Removed need for H-score for Tocilizumab use; added comment on very uncertain benefit and high risk of adverse events
V4	Jun 19, 2020	<ul style="list-style-type: none"> • Dexamethasone added for moderate-severe disease (moderate defined as needing supplemental oxygen; severe defined as needing mechanical ventilation) • Recommendation to NOT start antibiotics UNLESS strong suspicion of bacterial superinfection, or unless critically ill • Documentation of HScore required for Tocilizumab use • Added clinical trial contact info for outpatients and for prophylactic vs therapeutic LMWH in COVID-19 • Removed warning to avoid NSAIDs
V3	Apr 10, 2020	<ul style="list-style-type: none"> • Clinical trials for COVID initiated (trial coordinator x 32537) • Strong recommendation to D/C all antibiotics if no bacterial growth after 48 hours, and no clinical deterioration; • HScore (for sHLH) added to guide potential use of Tocilizumab (need HScore > 169 and refractory ARDS and ID/ICU approval) • No evidence for <i>therapeutic</i> anticoagulation; prophylactic LMWH as per admission guidelines • Removal of empiric oseltamivir for inpatients
V2	Mar 27, 2020	<ul style="list-style-type: none"> • Change in case definition • Removal of HCQ as standard of care for hospitalized patients <i>Rationale: early clinical experience at MUHC shows inpatients at MUHC are at high risk of arrhythmias but telemetry/code blue protocol logistically difficult on COVID wards; weak clinical evidence of efficacy of HCQ; upcoming RCT for COVID (info was N/A for earlier version) → risks now outweigh potential benefits.</i> • Doxycycline OR azithromycin (instead of azithro) for patients with CXR infiltrate • Tocilizumab <i>may</i> be considered for severe hyperinflammatory ARDS (HLH/CRS) in conjunction with ID/ICU • Reference to Admission guide for hospitalized patients

V1.2	Mar 20, 2020	<ul style="list-style-type: none">• Removal of oseltamivir prescription for <i>non-hospitalized</i> patients• Addition of comment that systemic steroids not indicated for COVID-19• Addition of comment that insufficient data to recommend adding or stopping ACE-inhibitors/ARBs
V1.1	Mar 19, 2020	<ul style="list-style-type: none">• Change in case definition (now any ILI regardless of travel or contact)• Changed recommendations re. quarantine: refer to IPC and Sante Publique
V1.0	March 18, 2020	Original version
