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The Potential Use of Hypochlorous Acid and a Smart Prefabricated Sanitising Chamber to Reduce Occupation-Related COVID-19 Exposure

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Abstract: This work is part of a project on the development of a smart prefabricated sanitising chamber (SPSC) to provide extra measures against the transmission of the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2). Stabilised hypochlorous acid (HOCl) is an approved disinfectant against SARS-CoV-2 by the Environmental Protection Association US in its liquid form on non-porous surfaces. This review is extended to cover its viricidal/bactericidal efficacy in aerosolised or sprayed form which showed an effective dose of as low as 20 ppm and the exposure duration of at least 60 s. The aerosolised application was also recommended with particle size of less than 200 μ m to increase the contact with pathogens. The review also includes the safety and toxicity of HOCl with different concentrations. The review calls for more investigations into the effect of HOCl in mist and fog form on the respiratory system when transitioning through the proposed SPSC. **Keywords:** SARS-CoV-2, Sanitiser, occupational health, cross-contamination, healthcare management

Introduction

The novel coronavirus, named severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2; COVID-19) has spread globally at an unprecedented rate with over 33 million infected cases and 1,012,341 deaths as of October 1, 2020.¹ In Australia, a suppression strategy has been adopted since the start of the pandemic with the aim of maintaining a low transmission rate close to 1 or below to make sure the healthcare system does not exceed its capacity and proper medical treatment is performed on infected patients.^{2,3}

To date, the extent of SARS-CoV-2 transmission and risk factors associated with infection are not fully understood; however, a scientific brief conducted by the World Health Organization (WHO) has stated that the main modes of transmission for SARS-CoV-2 include contact, droplet, airborne, fomite, fecal-oral and blood-borne with the potential for transmission between humans, mother-to-child and animal-to-human.^{4,5}

While strategies with social distancing including different levels of lockdown, mask use and increased personal hygiene are effective in slowing the COVID-19 transmission rate, recent studies have provided evidence on transmission via aerosol^{6,7} or the fecal shedding of the virus, making toilets significant objects of transmission as they aerosolise some of the contaminants within faeces.⁸ This indicates the need for

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further measures to effectively suppress the virus. Individual respiratory protective devices including N95 respirators and medical masks have been used as critical tools to protect healthcare workers against COVID-19.9 However, studies have found increased levels of discomfort, including higher facial skin temperature and adverse skin reactions, and these could induce non-compliance.^{10,11} As SARS-CoV-2 is known to remain viable for hours in aerosols and for days on surfaces,⁶ the importance of containment and prevention of cross-contamination in facilities with a high number of people frequently in contact with COVID-19 patients or people with high risk of carrying SARS-CoV-2 has been emphasised. Incorrect removal of personal protective equipment was known to contaminate 39% of healthcare workers with highly resistant micro-organisms.12 These containment measures are critical to public health safety especially where movement restrictions are relaxed, and new waves of infection are observed.¹³ These groups including doctors, nurses, aged-care workers, hospital staff or quarantine personnel. In the second wave of COVID-19 in Victoria, over 1,400 infected patients were healthcare workers and 70-80% of them were infected at work (as of August 27, 2020).¹⁴ This alarming infection rate not only puts the staff's health at risk but also hinders the response of the health service as trained professionals are isolated instead of giving care to patients. Particularly, more than 600 staff at Frankston Hospital, Victoria were in isolation following an outbreak at the facility.¹⁵

We have initiated a project on the development of a smart prefabricated sanitising chamber (SPSC) by way of aerosolised sanitation to disinfect COVID-19 exposed workers with minimal interruption of their operation and prevent cross-contamination. By minimising potential spread of SARS-CoV-2 at critical facilities such as COVID-19 treatment clinics and quarantine buildings for returned travellers, the use of the smart prefabricated sanitising chamber will help to prevent transmission into the broader community. Finding a suitable sanitiser is crucial to achieve a practical design for a SPSC (Figure 1).

A Potential Disinfectant

Wickramatillake and Kurukularatne¹⁶ analysed the viricidal efficacy of chloroxylenol, benzalkonium chloride and common bleaches against SARS-CoV-2, showing their effectiveness after 5 minutes of treatment. Hydrogen peroxide was also found to be used as a disinfectant against COVID-19 but this disinfectant has high reactivity and can act as a powerful oxidiser with metals.¹⁶ However, no commercially available data were shown on the efficacy of these disinfectants in aerosolised form or potential health impacts related to inhalation or contact with the skin. We have reviewed the evidence on the efficacy of hypochlorous acid (HOCl) in different settings via different applications and compared bactericidal and viricidal activity of hypochlorous acid to other disinfectants such as sodium hypochlorite (NaOCl), the most widely used disinfectant in the food industry, at the same free chlorine concentration.

HOCl is a weak acid which interacts with structural proteins such as the capsid or surface compounds, lipid envelope and DNA/RNA materials of viruses.¹⁷ The stability of HOCl is preserved by maintaining an optimal pH range of 3.5–5.5. HOCl can be generated by one of three methods: hydrolysis of chlorine gas, electrolysis of salt water, and acidification of hypochlorite. HOCl can be readily sourced commercially or generated inhouse. HOCl is listed in List N: Disinfectants for Use Against

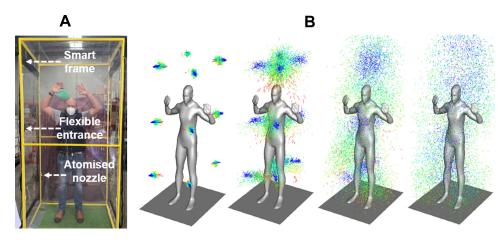


Figure I Prefabricated sanitising chamber: (A) prototype and (B) demonstration of aerosolised hypochlorous acid in the chamber.

EPA Registration Number	To Kill SARS-CoV-2, Follow Disinfection Directions For the Following Virus(es)	Contact Time (Minutes)	Product Name	Supplier
93908–1	Norovirus	10	Envirolyte O & G	Aqua Engineered Solution Inc.
92449–1	Rhinovirus	10	Annihilyte- I	Annihilare Medical Systems Inc.
87518–1	Norovirus	I	Hsp2O	HSP USA LLC
92108–1	Norovirus	10	Excelyte Vet	PCT Ltd
85134-1	Norovirus	10	Envirocleanse A	Envirocleanse LLC
91582–1	Adenovirus; Rhinovirus	10	Danolyte	Danolyte Global Inc.
89896–2	Human coronavirus	10	Cleansmart	Simple Science Ltd

Table I Currently Listed Hypochlorous Acid Products Approved by United States Environmental Protection Agency (EPA) for	or Use
Against SARS-CoV-2 as of October 1, 2020	

Note:Reproduced from United States Environmental Protection Agency [homepage on the Internet], US. 2020 August 20 List N: disinfectants for Use Against SARS-CoV-2 (COVID-19). Available from: https://www.epa.gov/pesticide-registration/list-n-disinfectants-use-against-sars-cov-2-covid-19.

SARS-CoV-2 by the US Environmental Protection Agency (as shown in Table 1).¹⁸

Disinfectants and disinfection techniques should be safe for a long exposure time on personnel, equipment and surfaces used in the fight against coronavirus. Disinfection technologies for indoors are important to reduce transmission, especially in crowded settings. HOCl has been proposed as an effective disinfectant in various settings from food production, handling and preparation (farming and hospitality industry) to healthcare applications, including chronic wound care and eyelid infection management, as listed Table 2.^{19–21}

Efficacy

While most disinfectants are tested or studied in their liquid form on non-porous surfaces, the viricidal/bactericidal efficacy of HOCl has also been studied when used in aerosolised/sprayed form and on porous surfaces. Hakimullah et al.¹⁷ present a comprehensive insight into the efficacy of HOCl in spray form to inactivate *Escherichia coli* and *Salmonella* with an effective concentration of 50–100 ppm on rayon fabric. Another study²² found that HOCl exhibited higher bactericidal activity (4.4 log₁₀ CFU/mL) compared with NaOCl (1.3 log₁₀CFU/mL) with the same concentration of 0.5 mg/L. Results from spray experiments¹² also showed that 20-minute exposure to sprayed HOCl of 0.01% was sufficient to reduce 99.5% of *Staphylococcus epidermidis*.

Another study by Wang et al.²³ found the potential pharmaceutical application of stabilised HOCl in the

control of soft tissue infection with a kill time of less than 2 minutes for 19 out of 20 pathogens. HOCl also had a shorter kill time than NaOCl and H₂O₂. Hao et al.²⁴ recommended the use of HOCl as a mouth wash and hand sanitiser (at 100-200 ppm) and for surface application. They also recommended spray or fog application with an aerosol size of less than 20 µm, due to the fact that smaller particles in spray form may help HOCl molecules to be suspended in the air for a longer duration. This low settling velocity rate may increase the solution's chance of coming into contact with pathogens and inactivating them. When the aerosol was not sprayed directly onto an inoculated surface, a lower amount of solution had a chance to come into contact with the avian influenza virus. It required at least 10 minutes of contact to be effective.²⁴

Another work²⁵ also found the effective concentration of HOCl to be 200 ppm in decontaminating inert surfaces carrying noroviruses and other enteric viruses in a 1-minute contact time. When diluted 10-fold, HOCl solutions at 20 ppm were still effective in decontaminating environmental surfaces carrying viruses in a 10-minute contact time.

Disinfection in the form of an effective surface disinfectant, hand hygiene or air disinfection, is the major preventive measure in containing the novel coronavirus. High-risk exposed surface areas need to be cleaned frequently with a suitable disinfectant. The characteristic features of an ideal disinfectant are a low contact time with significant antiviral activity with it being safe for humans, the environment, surfaces and

Table 2 Current Disinfecting	Applications of	Hypochlorous Acid
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Applications	Reference		
Hygiene solution on ocular skin	Stroman et al. ²⁰		
Wound care agent and topical application	Wang et al., ²³ Rasmussen and Williams, ²⁹ Del Rosso and Bhatia ³⁰		
Mouth wash and hand sanitizer	Hao et al. ²⁴		
Biosecurity enhancement in poultry production	Hakim et al. ¹⁷		
Decontamination agent against Shiga toxin-producing Escherichia coli on fresh beef	Cap et al. ²¹		
Disinfecting dental unit water lines	Martin and Gallagher ¹⁹		
Cleaning and disinfecting GI endoscopic procedure rooms	Overholt et al. ²⁶		
Mist application for enclosed area	Galvin et al. ²⁷		
General surface disinfectant	Hakim et al., ¹⁷ Hao et al. ²⁴ Rasmussen and Williams ²⁹ Block and Rowan ³¹		

Abbreviations: EPA, United States Environmental Protection Agency; FDA, US Food and Drug Administration; HOCI, hypochlorous acid; NaOCI, sodium hypochlorite; SARS-CoV-2, severe acute respiratory syndrome coronavirus 2; SPSC, smart prefabricated sanitising chamber; TGA, Australian Therapeutic Goods Administration; WHO, World Health Organization.

equipment. HOCl is approved by the Australian Therapeutic Goods Administration (TGA) as a disinfectant against COVID-19. The US Food and Drug Administration (FDA) has approved the use of hypochlorous acid for eye and dental spray application and as a no-rinse sanitiser for fruit and vegetables. With several studies on the sanitising power and effect of HOCl, it has been shown to eliminate viruses, bacteria and spores at an acceptable reduction log with effective concentration of as low as 20 ppm and a kill time as low as 1 minute. Further research is required to determine the optimum exposure time with non-direct contact spray HOCl to ensure the viricidal efficacy in settings such as the proposed SPSC.

Safety and Toxicity

In terms of the safety and toxicity owing to the potential direct contact with ocular, skin and respiratory systems, stabilised HOCl has been found to be non-irritating (rabbit eye) and non-sensitising (guinea pig) in animal models. No ocular irritation was observed when 0.013% HOCl was sprayed into the eyes of Dutch pigmented rabbits every 8 hours for 72 hours.²³ Guinea pigs showed no evidence of dermal reaction when concentrations of 0.01%, 0.03%, and 0.10% w/v stabilised HOCl were applied topically. A 28-day toxicity study with the same concentrations as the dermal study showed no evidence of systemic toxicity.²³ HOCl was also found to be an effective disinfectant that was 80–200 times more

effective than standard disinfecting procedures while being non-toxic to humans.²⁶ Van Doremalen et al.⁶ and Galvin et al.²⁷ recommended HOCl use for offices with an emphasis on its use for dental clinics and the importance of fog or mist application²⁸ of the disinfection given the aerosol property of the virus.

HOCl is safe and non-toxic for human and environmental use. It is also inexpensive with the flexibility of applying methods (spray mist, fog, liquid) which makes it an appropriate candidate as a choice of disinfectant for indoor settings on a regular basis. The presence of organic and reducing agents significantly reduces the effectiveness of hypochlorous acid²¹ and this is an area that needs to be studied further to provide robust conclusions. This review recommends further study in regard to the safety of HOCl with mist and fog application on respiratory system, skin and eyes, in the smart prefabricated sanitising chamber setting, acknowledging that the deployment of Brio HOCL[™] inactivates viruses, bacteria, endospores, and fungi and is safe for human tissues (including eye, lung, and skin).²⁹ It should be noted that mist applications have already been approved for eyelid infection management,²⁰ skin²³ and cosmetics³⁰ with repeated daily exposure for a set period of time. Other reviews by Block and Rowan³¹ and Scarano et al.³² found that hypochlorous acid had high predictability and low cost to disinfect against COVID-19 in oral-maxillofacial clinics by spraying small aerosol

particles. This review calls for more investigations into the effect of HOCl in mist and fog form on the skin, eyes, and respiratory system when exposed in the proposed smart prefabricated sanitising chamber, especially in the presence of organic substances.

Conclusion

In this review on the use of stabilised hypochlorous acid (HOCl) as a potential disinfecting agent for the smart prefabricated sanitising chamber application, the current evidence on the viricidal efficacy, safety, and toxicity are presented. HOCl stabilised at the pH of 3.5-5.5 is a weak acid which interacts with structural proteins or viral materials to inactivate micro-organisms. HOCl is currently a disinfectant for SARS-CoV-2 approved by the US Environmental Protection Association under different brands. HOCl with a concentration as low as 20 ppm was found to be effective in disinfecting surfaces including porous rayon. Aerosolised or sprayed HOCl is effective in eliminating micro-strains such as Staphylococcus epidermidis after an exposure of 1 min to 20 mg/L (equivalent to 20 ppm). The safety and non-toxicity for humans and the environment, the low cost, and the flexibility of applying methods make HOCl an appropriate candidate of disinfectant for indoor settings such as offices, hospitals, and healthcare clinics. Further investigations into the effect of HOCl in mist and fog form on the skin, eyes, and respiratory system especially in special settings such as the proposed smart prefabricated sanitising chamber with presence of organic materials is recommended.

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Disclosure

The authors report no conflicts of interest in this work.

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