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ultrasound assessment for the surveillance of lung health

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Abstract

Background: SARS-CoV-2 infection, manifesting as COVID-19 pneumonia, constitutes a global pandemic that is disrupting health-care systems. Most patients who are infected are asymptomatic/pauci-symptomatic can safely self-isolate at home. However, even previously healthy individuals can deteriorate rapidly with life-threatening respiratory failure characterized by disproportionate hypoxemic failure compared to symptoms. Ultrasound findings have been proposed as an early indicator of progression to severe disease. Furthermore, ultrasound is a safe imaging modality that can be performed by novice users remotely guided by experts. We thus examined the feasibility of utilizing common household informatic-technologies to facilitate self-performed lung ultrasound.

Methods: A lung ultrasound expert remotely mentored and guided participants to image their own chests with a hand-held ultrasound transducer. The results were evaluated in real time by the mentor, and independently scored by three independent experts [planned a priori]. The primary outcomes were feasibility in obtaining good-quality interpretable images from each anatomic location recommended for COVID-19 diagnosis.

Results: Twenty-seven adults volunteered. All could be guided to obtain images of the pleura of the 8 anterior and lateral lung zones (216/216 attempts). These images were rated as interpretable by the 3 experts in 99.8% (647/648) of reviews. Fully imaging one's posterior region was harder; only 108/162 (66%) of image acquisitions was possible. Of these, 99.3% of images were interpretable in blinded evaluations. However, 52/54 (96%) of participants could image their lower posterior lung bases, where COVID-19 is most common, with 99.3% rated as interpretable.

Conclusions: Ultrasound-novice adults at risk for COVID-19 deterioration can be successfully mentored using freely available software and low-cost ultrasound devices to provide meaningful lung ultrasound surveillance of themselves that could potentially stratify asymptomatic/paucisymptomatic patients with early risk factors for serious disease. Further studies examining practical logistics should be conducted.

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Keywords: COVID-19, Telemedicine, Lung Sonography, Remote Telementoring, Community Health, SARS-CoV-2, Patient focused health-care

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Introduction

COVID-19 pneumonia is disrupting life on the planet earth in an unprecedented fashion. While many if not most people have mild or asymptomatic disease, others, even young previously healthy people, may become

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rapidly sick with severe hypoxia despite exhibiting minimal symptoms, including dyspnea [1–3]. The burden on health care systems may be extraordinary, with even well-developed nations' health care systems being overwhelmed. Health care providers may be particularly susceptible if appropriate infection prevention and control measures are not in place [4]. Thus, solutions need to be sought to provide excellent patient care, but also to protect provider health. COVID-19 is a paradox, as despite the risk to providers, the majority contracting the virus will not develop COVID-19 pneumonia. Most will have none or minor symptoms and can safely self-isolate at home. However, those who develop severe disease, need to be identified early [4].

Compared to chest radiographs and computed tomography (CT), lung ultrasound (LUS) is a simpler, more portable, economical, and potentially home-based technology that might be used for at-risk patients to selfmonitor their lungs of for early signs of COVID-19 pneumonia [5–7]. Findings from COVID-19 pneumonia are typically present in the lung periphery [8-10], an anatomic fact that permits LUS to be used to diagnose and manage all phases of care in COVID-19 [5, 9, 11]. LUS may detect early disease progression as the lungs deteriorate from normal to an alveolar-interstitial pattern of lung disease with single discrete vertical artifacts (B-lines) or confluent B-lines [9, 11]. Through work onboard the International Space Station examining self-performed telementored lung ultrasonography (SPTMLUS) performed by inexperienced point-of-care users guided by remote experts [12], we have long known that accurate ultrasound images of the lungs can be self-obtained [13-17]. What has never been examined, is whether willing but ultrasound-naïve adults can be remotely mentored to obtain meaningful lung ultrasound images upon themselves to triage alveolar-interstitial pneumonic diseases, such as COVID-19. The purpose of this study was thus to examine the feasibility and quality of SPTMLUS of novices when expertly guided. Furthermore, this paradigm may be considered a specific example of a broader concept that may contribute to many facets of patient focused and individualized healthcare.

Methods

This study was registered and ethically approved and structured to comply with the SQUIRE reporting guidelines [18]. A healthy cohort of self-isolating participants in Edmonton, Alberta conducted SPTMLUS examinations mentored by a remote expert. The participants were self-isolating among their family units in response to applicable Public Health orders in effect. After informed consent, participants completed an electronic demographic survey (Additional file 1) and received a package containing a disinfected hand-held high-frequency linear ultrasound probe (Philips Lumify, Philips, Amsterdam, NL), and a package of sterile ultrasound-gel. They also watched a brief instructional video on how to hold the probe and where anatomically they would be guided to scan (Additional file 2).

Thereafter, a lung ultrasound expert in Calgary (AWK) guided the subjects to measure their blood pressure (details reported elsewhere) and conduct a standardized lung examination, using Zoom Teleconferencing (Zoom, Hillsboro, OH). The desired examination was based on the 14-zone method proposed by Soldati for International Standardization of the Use of Lung Ultrasound for Patients with COVID-19 [19] (Fig. 1). The study goal was to generate an adequate "Batwing" depiction of the pleura interface between two rib shadows at each location on the thorax [20] (Fig. 2). Outcome measures were, therefore, (1) whether the subject was physically able to reach all 14 desired anatomic locations (Figs. 3 and 4) and (2) whether the quality of images was considered "adequate" for image interpretation and diagnosis. Images were optimized through remote control of ultrasound "knobology" by the mentor using remote access software (Teamviwer, Göppingen, Germany). The pleural interface was interrogated with 2D, M-mode, and color-Power Doppler (CPD) modes[21], and all image acquisition attempts were videorecorded. The mentor scored each of the pleural images real time using the proposed Soldati method from 0 (normal) to 3 (very abnormal) (Additional file 4) [19], and counted the number of B-lines present at each anatomic location. Each participant completed an online post-test evaluation that included their perceptions of the difficulty in performing their self-examination including a 5-point Likert scale rating the examination at each location as being one of; 1-Very Hard, 2-Hard, 3-Neutral, 4—Easy, 5—Very Easy (Additional file 3).

Subsequently the mentor retrospectively reviewed all examinations and prepared representative still images and videos at each anatomic location, which were uploaded to a protected website for an a-priori planned independent review by three outside lung ultrasound experts, who rated both the image quality (as adequate vs. inadequate) and the degree of abnormality using the same Soldati scoring protocol[22]. If they had any concerns regarding image quality, they were encouraged to review the entire source examination. Standard descriptive statistics were used. Patients reporting and not reporting any upper body musculoskeletal concerns or shoulder injuries were analyzed separately and the comparisons between groups were made using Fisher's exact tests. The theoretical ability to complete commonly used and recommended lung ultrasound imaging protocols was calculated by analyzing the ability for SPTMLUS





Fig. 2 Representative still image of mentee self-performing lung ultrasound to demonstrate the pleural interface. Ultrasound Naïve participant being instructed to demonstrate the pleural interface of the visceral and parietal pleural illustrating the "Batwing" sign of Lichtenstein [20]. It should be noted that all lung ultrasound is a dynamic examination better viewed real time and in video recordings (Additional file 4) than still images



Fig. 3 Representative still image of mentee self-performing lung ultrasound to demonstrate the pleural interface of her back. Ultrasound Naive participant being instructed to demonstrate the pleural interface of the visceral and parietal pleural illustrating the "Seahorse" sign of Lichtenstein [20]. It should be noted that all lung ultrasound is a dynamic examination better viewed real time and in video recordings (Additional file 3) than still images





to be completed at each recommended location of the Extended Focused Assessment with Sonography for Trauma [23], the Bedside Lung Ultrasound Assessment [24], and the International evidence-based recommendation for point-of-care lung ultrasound [25], and a theoretical examination just looking at the anterior, lateral, and posterior lung bases (Fig. 5).

Results

Demographics

The study population consisted of a convenience sample of 27 self-isolating inhabitants in Edmonton, Alberta, Canada. All participants stated they were comfortable with typical informatic tools of modern society, such as smart-phones and tablet computers, but none were informatic technology experts. Three [11.1%] had prior ultrasound exposure but none had ever performed LUS (Table 1). Fifteen [14.8%)



the public domain, all the data available to the original reviewers is not available here due to confidentiality considerations. Videorecording of the complete examinations for those subjects who agree to disclose their personal images are available in Additional file 3

of participants had a Masters or higher Education degree, 7 [26%] reported an upper body musculoskeletal problem and 8 [29.6%] reported a previous shoulder injury (Table 1).

Ability to complete a comprehensive pleural ultrasound examination

Overall, all 27 participants were able to obtain images from all (100%) of the eight anatomic locations on the anterolateral chest. The back was relatively harder, although possible on the lower back in 52/54 (96.3%) of attempts; 38/54 (70.4%) midback, and only18/54 (33%) on the upper back (Table 2). When asked the subjective difficulty in performing the examination, participants rated imaging their own anterior and lateral chest as between "easy" and "very easy" in all locations, while the back was rated between "neutral" to

 Table 1
 Demographic profile of the 27 ultrasound-novice selfisolating participants

Demographics	Response (standard deviation) [percentage]	
Age	42.81 years (2.33)	
Male/Female	15 [56%]/12 [44%]	
Height	172.9 cm (10.20)	
Weight	80.67 kg (17.82)	
BMI	27.0	
Highest level of education		
Did not complete High School	2 [7.4%]	
High school	4 [14.8%]	
Diploma after high school	8 [29.6%]	
Trade certificate	1 [3.7%]	
Undergraduate degree	7 [25.9%]	
Master's degree	4 [14.8%]	
Other	1 [3.7%]	
Ever held an ultrasound?		
Regularly	0 [0.0%]	
Once ¹	3 [11.1%]	
Never	24 [88.9%]	
Pre-existing respiratory concerns?		
Yes ²	4 [14.8%]	
No	23 [85.2%]	
Pre-existing cardiac concerns		
Cardiac bypass	2 [7.4%]	
Premature atrial contractions	1 [3.7%]	
None	24 [88.9%]	
Smoking or vaping		
Cannabis only	2 [7.4%]	
Tobacco only	1 [3.7%]	
None	24 [88.9%]	
Upper body musculoskeletal problems		
None	20 [74.1%]	
Present	7 [25.9%]	
History of shoulder injury		
Yes	8 [29.6%]	
No	19 [70.4%]	
Asthma diagnosis		
Yes	4 [14.8%]	
No	23 [85.2%]	

¹ Three subjects replied affirmative, none had formal training; one had unofficially been shown Obstetrical ultrasound in Uganda, one Firefighter had previously participated in just-in-time mentored abdominal and vascular ultrasound; one research coordinator had been exposed to an ultrasound phantom: ²all reported pre-existing respiratory concerns were self-reported asthma

"easy" in all locations (Table 3). There was no statistical relation between the ability/inability to obtain images

and a history of shoulder injury at any of the anatomic location.

Real-time subjective mentored evaluation and a-priori independent review

Although not all desired locations in all participants could be imaged, of the sites that could, the image quality of 322/324 (99.7%) of real-time determinations was felt to be adequate in real time assessment and adequate in 1124/1128 (99.6%) of independent reviews (Table 2). An examination of suspected reasons for inadequate images is presented in Table 4. Results regarding evaluated lung score and B-line counts are available (Additional file 5), recognizing that no patient had known COVID-19 during examination.

Selected standardized lung ultrasound performance difficulty

The theoretical feasibility and participant-rated ease of performance of commonly used LUS protocols is presented in Table 5. The Extended Focused Assessment with Sonography for Trauma (EFAST) examination [23] was 100% feasible and would be "very easy" (mean Likert score 4.67 out of 5) as assessed by the participants, as was the Bedside Lung Ultrasound in Emergency (BLUE), (feasibility 100%; ease of performance score 4.56) [20]. Similarly, the International evidence-based recommendations for point-of care lung ultrasound (ICC-LUS) was 100% feasible and easy (mean score 4.56) [26]. Even the overall Soldati COVID examination was feasible in 85.7% of locations and overall rated easy (mean score 4.07). A new theoretical examination involving the lowermost lung fields of the anterior, lateral, and posterior lung bases would be 98.8% feasible and easy to very easy (mean score 4.37) (Table 5).

Discussion

Our results demonstrate that adults without prior ultrasound experience were remarkably adept at accurately imaging their own chests to generate clinically meaningful images under the guidance of a remote mentor. Every participant was able to satisfactorily image their anterior, lateral, and lower posterior thoracic areas. With the exception of one participant who later contracted COVID-19 [27], all participants were at risk, but remained clinically well. In our opinion, the most important interpretation of these results, beyond the implications to assist with the surveillance of COVID-19, is the potential to explore the paradigm of mentored self-care to other conditions involving remote, self-isolated, or vulnerable populations to better enable and empower their own health maintenance. Furthermore, although the technique required an economical hand-held ultrasound

Anatomic Location	Image generation deemed possible ^a	lmage quality deemed adequate (real time mentor) ^b	lmage quality deemed adequate (independent review) ^c
(1) Right upper chest	100%	100%	100%
(2) Right lower chest	100%	100%	100%
(3) Left upper chest	100%	100%	100%
(4) Left lower chest	100%	100%	98.8%
(5) Right side lower	100%	100%	100%
(6) Right side upper	100%	100%	100%
(7) Left side lower	100%	100%	100%
(8) Left side upper	100%	100%	100%
(9) Right back lower	100%	100%	98.7%
(10) Right back middle	74.1%	95.2%	98.4%
(11) Right back upper	70.4%	100%	100%
(12) Left back lower	100%	100%	100%
(15) Left back middle	66.7%	100%	100%
(16) Left back upper	37%	100%	100%

Table 2 Aggregate ability to obtain images from the complete thorax and the assessed quality of the images that could be obtained

Anatomic location is visually demonstrated in Fig. 1

^a Image Generation was the remote mentoring expert's assessment of the interaction with the participant as to whether assessing the anatomic location was possible to image

^b Image Quality was assessed real time by the Mentor during the examination

^c Image Quality was assessed as the composite result of the a-prior independent review of the images/videos

Table 3 Participants subjective post-test assessment of difficulty in providing self-administered ultrasound examination

Anatomic	Median	Mean Self-Reported Score		
Location	Rating	(95% confidence interval)		
(1) Right Upper Chest	5	4.67 (4.24–5.09)		
(2) Right Lower Chest	5	4.67 (4.24–5.09)		
(3) Left Upper Chest	5	4.67 (4.24–5.09)		
(4) Left Lower Chest	5	4.67 (4.24–5.09)		
(5) Right Side Lower	5	4.52 (4.07-4.96)		
(6) Right Side Upper	5	4.44 (4.04-4.85)		
(7) Left Side Lower	5	4.52 (4.07–4.96)		
(8) Left Side Upper	5	4.44 (4.04–4.85)		
(9) Right Back Lower	4	3.89 (3.49-4.29)		
(10) Right Back Middle	3.5	3.41 (3.01-3.81)		
(11) Right Back Upper	3	2.93 (2.41-3.44)		
(12) Left Back Lower	4	3.96 (3.55-4.38)		
(15) Left Back Middle	4	3.35 (2.89–3.80)		
(16) Left Back Upper	3.5	2.85 (2.33-3.37)		

Anatomic location is visually demonstrated in Fig. 1

Mean Self-Reported Score¹ was recorded as a 5-point Likert scale from 1 Hard to 5 Easy (1—Very Hard, 2—Hard, 3—Neutral, 4—Easy, 5—Very Easy)

device, all other communication technologies are widely available in many if not most homes currently, thus minimizing costs and maximizing potential opportunities.

Challenges related to COVID-19 include the high infectivity of the virus, rapid mutations of more aggressive variants, the frequency of asymptomatic carriers, and the fact that pauci-symptomatic patients may shed the most virus immediately before exhibiting any symptoms. This makes in-person medical assessments a potentially dangerous undertaking, and one that contributed to the near collapse of many healthcare systems [28, 29]. Furthermore, COVID-19 is predicted to be just one of many future zoonotic-based pandemics that will afflict humans in the future [30]. Thus, early experiences with nearly overwhelmed health systems, prompted recommendations to employ telemedical capabilities to provide advanced outreach capabilities, for the "entire population not only for hospitals" [31]. Such an approach would hospitalize only those with severe disease, with asymptomatic or paucisymptomatic observed at home, thereby decreasing contagion and preserving personal protective equipment [8, 31]. Such an approach necessitates the ability to quickly recognize those who exhibiting signs of deterioration, and "rescuing" them quickly [1, 8]. Previous work has demonstrated that with remote guidance, non-expert point of care providers, who may be as inexperienced as children, can be guided to place an inexpensive ultrasound probe onto the chest to assess the visceral-parietal pleural surface [17].

LUS, a relatively new discipline based on the science of artifact analysis, is now established in emergency and

Tak	blo	e 4	Suspected	d reasons :	for inad	lequate c	qualit	y images

Anatomic Location	Reason for Image Inadequacy		
Left lower anterior chest	Previous median sternotomy with pericardiotomy and cardiac surgery		
Right lower back	Technical issue where patient video image obscured the ultrasound image on the recorded video		
Right middle back	Technical issue where patient video image obscured the ultrasound image on the recorded video		

Table 5 Average naïve mentee scores for selected lung ultrasound protocol performance feasibility and difficulty

Lung ultrasound protocol	Anatomic locations	Feasibility	Average self-rated naïve user score
EFAST examination	1,2,3,4	108/108 (100%)	4.67 (Very Easy)
BLUE examination	1,2,3,4,5,6,7,8	216/216 (100%)	4.56 (Very Easy)
Soldati COVID Examination	1,2,3,4,5,6,7,8, 9,10,11,12,15, 16	324/378 (85.7%)	4.07 (Easy)
ICC-LUS Examination	1,2,3,4,5,6,7,8	216/216 (100%)	4.56 (Very Easy)
Lower Lung Fields	2,4,5,7,9,12	160/162 (98.8%)	4.37 (Easy—Very Easy)

EFAST Extended Focused Assessment with Sonography for Trauma [23], *BLUE* Bedside Lung Ultrasound Assessment[44], *Soldati COVID examination* Proposal for International Standardization of the Use of Lung Ultrasound for Patients With COVID-19[22], *ICC-LUS Examination* International evidence-based recommendations for point-of-care lung ultrasound[26]. Mean Self-Reported Score was recorded as a 5-point Likert scale from 1 Hard to 5 Easy (1—Very Hard, 2—Hard, 3—Neutral, 4—Easy, 5—Very Easy)

resuscitative medicine [26]. We and others recognize its near unlimited value to manage the COVID-19 crisis [11, 19, 32]. The gold standard diagnostic test for COVID-19 diagnosis is the identification of viral nucleic acids (PCR). However, compared to PCR, CT scan may show disease at an earlier time frame [33], as may LUS [34], even in cases, where the PCR was initially negative [35]. As lung ultrasound is a technology that may diagnosis COVID-19 earlier than PCR testing, is portable and able to go to the patient, relatively economical, can discern the presence of progression of COVID-19 pneumonia, and is easily repeatable over the projected time course of disease, lung ultrasound may be beneficial to follow pleural health over the complete evolution of disease [11]. Through marrying the ability to perform ultrasound upon oneself using remote guidance, all these potential uses might be provided without ever requiring a physical encounter with a health-care provider to completely reduce health care exposure to infection during the initial assessment and risk stratification of healthy but at-risk individuals.

In health, the pleural interface of the normal lung of a healthy human, is typically the only part of the lung that can be viewed with ultrasound [36], as air has the highest acoustic density of any other component of the human body, effectively preventing transmission of ultrasound waves beyond the pleura. Thus, when examining the interface of the parietal pleura of the chest wall and the visceral pleura of the lung, only this interface can be seen. When the lung becomes diseased with COVID pneumonia it has been observed that abnormal lung artifacts arise at the pleural junction [11]. Of note, the patchy bilateral, multifocal ground-glass opacity and abnormalities typically found on CT associated with COVID-19 are predominantly identified in the lower [9, 37, 38] and posterior lung zones [9, 39], the so called COVID hot spots. At the bedside, a critical "tipping-point" for concern stratifying those with home-manageable minor diseases from potentially severe deterioration may be an evolution from the normal A-line pattern of health to a progressively abnormal alveolar-interstitial pattern of lung disease illustrated by initially single or discernable B line artifacts, progressing to confluent vertical artifacts (B-lines) or confluent vertical artifacts, ultimately culminating with white lungs and/or consolidated effusions of severe COVID-19 [11].

Although standard examinations have been proposed [9, 22], there is no single currently accepted "standard" examination. It was not unexpected that participants would be less adept at examining their backs, and a posterior examination does not comprise part of many common lung ultrasound protocols (Table 5). Nonetheless, 96.3% of all participants could image their own lower back, higher than was originally expected, and emphasizing the remarkable abilities of motivated laypersons when mentored.

COVID-19 may be particularly treacherous as there may be profound disassociation between the severity of hypoxemia and preservation of respiratory muscle mechanics, lung compliance, and an absence of dyspnea [40]. Thus, there are patients with COVID-19 who

exhibit oxygen levels incompatible with life without dyspnea, sometimes termed "happy hypoxia" but is more precisely termed silent hypoxemia [2]. There are also more than a few limitations of pulse oximetry such as O₂ Saturation monitoring may not be accurate at very low PaO₂ [2]. This condition may have an alarming frequency. Busana noted that among patients presenting to hospital with hypoxia consistent with acute respiratory failure, one-third were not dyspneic, including 18% with severely abnormal PaO_{2/}FiO₂ ratios of between 50 and 150, and overall, these patients had a mortality rate of 17.6% [1]. Thus, lung ultrasound offers a technique to potentially detect these patients at the earliest signs of lung swelling. Opportunities for future research include formal statistical comparisons of test performance characteristics of each of these modalities individually. However, we suspect that in actual clinical practice, these modalities would be complementary when used together. Recent guidelines for the use of Lung Ultrasound in managing COVID-19 did NOT recommend serial LUS re-examinations despite its potential utility in managing the patient's status due to concerns regarding infection transmission risks [9]. However, with the technique of RTMSPLUS these risks are completely obviated. Furthermore, these same guidelines, recommend that LUS should, however, be the initial lung imaging modality of choice in patients with minimal symptoms as LUS has higher sensitivity and lower radiation risks [9]. We have also demonstrated unmanned arial vehicle (drone) delivery of RTMSPLUS, with unlimited potential to access remote and disadvantaged populations [41].

Pneumothoraces, posterior lung fields, and the lung bases

Although the additional modalities of M-Mode and CPD are not necessarily critical to the inference of COVID-19, they contribute to the diagnosis of pneumothorax which recent guidelines recommend assessing for during a COVID-19 lung ultrasound examination [9]. Recent guidelines also strongly recommend that in addition to the usual anterolateral lungs, that posterior lung zones should be scanned whenever possible [9]. This is an aspect, wherein the healthier asymptomatic/Paucisymptomatic patients are advantaged in not being bedbound and supine and were able to image their lower posterior lung fields 100% of the time.

A potential limitation of our study was that our participants were not sick and were feeling well. Therefore, the majority of the LUS images were normal. However, this is appropriate for a screening test intended for asymptomatic/pauci-symptomatic populations. Our participants were relatively young and nearly all completed high school. Thus, our results may not generalize to an older, or less educated population. Our participants all had access to a computer and internet. Remote guidance may be more challenging in situations, where internet is not available, although we agree that internet availability should be a basic human right [42]. Three participants had previous ultrasound exposure, but none had examined the pleura before. In general males were less able to image their complete backs than females, but this may be less critical for lung surveillance in COVID-19, given the propensity for early COVID to affect the lung bases. Therefore, it may be more important for RTMSPLUS to examine the lung bases (anterior, lateral, and posterior locations) and in our study, the ability to image these locations was 100% (Table 2).

This study constitutes a proof of concept that we hope generates further discussion and analysis of the capabilities, logistical requirements, and human factors challenges in assisting in remotely mentored self-care. Clearly innumerable details require study before such a paradigm is ready for clinical application. Given these practicalities, however, we contend that the concept potentially empowers those remote from fixed hospital care to empower their own healthcare through permitting required imaging of their own anatomy and physiology wherever they are geographically as long as they are connected. Just as necessity drove space medicine to consider innumerable innovative ways to incorporate remotely mentored ultrasound into care paradigms [43], just-in-time mentoring of the isolated may have innumerable applications that deserve examination.

Conclusions

We contend that providing home-SPRTLUS may be a useful method to provide surveillance of at-risk populations. Besides earlier diagnosis and rescue of severe cases, we postulate that such a proactive approach empowering the willing to manage their own health might also potentially reduce anxiety and increase the "connectedness" of patients and health care providers, an intangible commodity that is severely threatened in these times of strict self-isolation.

Supplementary Information

The online version contains supplementary material available at https://doi. org/10.1186/s13089-021-00250-6.

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Authors' contributions

AWK made substantial contributions to the conception AND design of the work; AND the acquisition, analysis, AND interpretation of data; AND drafted the work AND substantively revised it AND approved the submitted version AND agreed both to be personally accountable for the author's own contributions and to ensure that questions related to the accuracy or integrity of any part of the work, even ones in which the author was not personally involved, are appropriately investigated, resolved, and the resolution documented in the literature. JLM made substantial contributions to the conception AND design of the work; AND the acquisition, analysis, AND interpretation of data; AND approved the submitted version AND agreed both to be personally accountable for the author's own contributions and to ensure that guestions related to the accuracy or integrity of any part of the work, even ones in which the author was not personally involved, are appropriately investigated, resolved, and the resolution documented in the literature. CGB made substantial contributions to the acquisition, analysis, AND interpretation of data; AND substantively revised the work AND approved the submitted version AND agreed both to be personally accountable for the author's own contributions and to ensure that questions related to the accuracy or integrity of any part of the work, even ones in which the author was not personally involved, are appropriately investigated, resolved, and the resolution documented in the literature. IWYM made substantial contributions to the acquisition, analysis, AND interpretation of data; AND substantively revised the work AND approved the submitted version AND agreed both to be personally accountable for the author's own contributions and to ensure that questions related to the accuracy or integrity of any part of the work, even ones in which the author was not personally involved, are appropriately investigated, resolved, and the resolution documented in the literature. LAM made substantial contributions to the acquisition, analysis, AND interpretation of data; AND substantively revised the work AND approved the submitted version AND agreed both to be personally accountable for the author's own contributions and to ensure that questions related to the accuracy or integrity of any part of the work, even ones in which the author was not personally involved, are appropriately investigated, resolved, and the resolution documented in the literature.

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Availability of data and materials

The data sets used and/or analysed during the current study are available from the corresponding author on reasonable request.

Declarations

Ethics approval and consent to participate

This research was approved by the CHREB of the University of Calgary, REB14-0634_MOD4.Consent for publication. All subjects provided written informed consent after full explanation of the study.

Consent for Publication

The research participants freely consent to publication of the main manuscript and Supplementary material.

Competing interests

Andrew W Kirkpatrick has consulted for the Zoll, Innovative Trauma Care, CSL Behring and SAM Medical Corporations, and is the PI of a Prospective Randomized Controlled Trial partially supported by the Acelity Corporation (https://clinicaltrials.gov/ct2/show/NCT03163095). He discloses a personal relationship with JL McKee. Jessica L McKee is the Research Director of Innovative Trauma Care and has consulted for Zoll, Aceso, SAM Medical Products and Acelity. She also discloses a personal relationship with AW Kirkpatrick. Chad G Ball MD, reported no disclosures. Irene W. Y. Ma, is a chair holder of the John A. Buchanan Chair in General Internal Medicine at the University of Calgary. She has no other conflicts of interest to disclose. Lawrence A. Melniker, reported no disclosures.

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