

Smartphone movement sensors for the remote monitoring of respiratory rates: Technical validation

Digital Health Volume 8: 1–11 © The Author(s) 2022 Article reuse guidelines: sagepub.com/journals-permissions DOI: 10.1177/20552076221089090 journals.sagepub.com/home/dhj



Sophie Valentine, Adam C. Cunningham, Benjamin Klasmer, Mohammad Dabbah, Marko Balabanovic, Mert Aral, Dan Vahdat and David Plans (D)

Abstract

Background: Mobile health (mHealth) offers potential benefits to both patients and healthcare systems. Existing remote technologies to measure respiratory rates have limitations such as cost, accessibility and reliability. Using smartphone sensors to measure respiratory rates may offer a potential solution to these issues.

Objective: The aim of this study was to conduct a comprehensive assessment of a novel mHealth smartphone application designed to measure respiratory rates using movement sensors.

Methods: In Study 1, 15 participants simultaneously measured their respiratory rates with the app and a Food and Drug Administration-cleared reference device. A novel reference analysis method to allow the app to be evaluated 'in the wild' was also developed. In Study 2, 165 participants measured their respiratory rates using the app, and these measures were compared to the novel reference. The usability of the app was also assessed in both studies.

Results: The app, when compared to the Food and Drug Administration-cleared and novel references, respectively, showed a mean absolute error of 1.65 (SD = 1.49) and 1.14 (1.44), relative mean absolute error of 12.2 (9.23) and 9.5 (18.70) and bias of 0.81 (limits of agreement = -3.27 to 4.89) and 0.08 (-3.68 to 3.51). Pearson correlation coefficients were 0.700 and 0.885. Ninety-three percent of participants successfully operated the app on their first use.

Conclusions: The accuracy and usability of the app demonstrated here in individuals with a normal respiratory rate range show promise for the use of mHealth solutions employing smartphone sensors to remotely monitor respiratory rates. Further research should validate the benefits that this technology may offer patients and healthcare systems.

Keywords

Smartphone, mobile phone, mHealth, digital health, mobile apps, remote monitoring, respiratory rate, gyroscope, sensors, accuracy, usability, software validation

Submission date: 29 November 2021; Acceptance date: 6 March 2022

Introduction

Extensive growth in the development and adoption of remote healthcare tools has been seen in recent years in response to the increasing demand for traditional offerings.¹ Notably, the COVID-19 pandemic has made salient how these mobile health (mHealth) tools may support healthcare systems to manage their patients when resources are pushed to breaking point.^{2–4} As more widely accessible tools can

be used by more people – and therefore offer greater impact – many mHealth smartphone applications (apps) have been developed due to the high global penetration of

Huma Therapeutics Ltd, London, UK Corresponding author:

David Plans, Huma Therapeutics Ltd, London, UK. Email: david.plans@psy.ox.ac.uk

Creative Commons Non Commercial CC BY-NC: This article is distributed under the terms of the Creative Commons Attribution-NonCommercial 4.0 License (https://creativecommons.org/licenses/by-nc/4.0/) which permits non-commercial use, reproduction and distribution of the work without further permission provided the original work is attributed as specified on the SAGE and Open Access page (https://us.sagepub.com/en-us/nam/ open-access-at-sage). smartphones. These systems offer a wide variety of services from telemedicine to remote monitoring and self-care, and evidence suggests they may produce improved economic⁵ and health outcomes.⁶

The respiratory rate (RR) is a fundamental indicator of health status for many health conditions, both general and specific to the respiratory system.⁷⁻¹² As such, mHealth solutions for monitoring of RR may offer significant value to patients and healthcare professionals (HCPs) alike. Although several hardware and software-based solutions exist to measure RR, currently they often have disadvantages. Hardware-based solutions, including piezoelectric sensors,¹³ pulse oximeters¹⁴ and multi-sensor devices,^{15,16} are typically expensive, vulnerable to limited means of manufacture and distribution¹⁷ and may lack interoperability with other health records, which is cited as a critical risk to the decentralisation of national healthcare systems.¹⁸ Software-based solutions address limitations of cost, manufacture and distribution; however, they typically employ less-stable mechanisms of action. These mHealth apps often use smartphone cameras or microphones,¹⁹⁻²¹ the latter of which have been evidenced to be vulnerable to environmental noise at the cost of accuracy and usability.20,22

Movement sensors may present a promising alternative software-based solution for mHealth RR monitoring. Research indicates that multi-axial accelerometers and gyroscopes – as found ubiquitously in modern smartphones – can accurately capture RR based on chest movements.^{23–30} Additionally, due to their mechanism of action, these sensors are significantly less affected by environmental noise. Overall, smartphone-based measurement of RR provides a potential low cost, and a widely available method for RR measurement, both in a remote monitoring environment and in locations where specialised hardware and software are not available.

This article presents the technical validation of a novel user-centric mHealth smartphone app that measures RR using the tri-axial gyroscope. We first conducted a preliminary evaluation of the device and study methods via a small lab-based study and then jointly assessed accuracy and usability on a greater scale and ecological valid environment via a remote study. Ethical approval was provided by the University of Exeter's Research Ethics Board (application ID: eUEBS004088), and all research was conducted in compliance with the Declaration of Helsinki.

Study 1

Methods

The preliminary evaluation pursued three aims: (a) to establish the accuracy of the novel mHealth smartphone app relative to a reference device cleared by the US Food and Drug Administration (FDA), (b) to understand the usability of the mHealth app and (c) to evaluate the suitability of a novel reference method that would permit accuracy assessments to be conducted via remote and real-world studies. Through a prospective, non-interventional, non-randomised study conducted on healthy volunteers, RR estimates provided by the FDA-cleared reference device were compared to those from the novel mHealth smartphone app and the novel reference. RR measurements were recorded simultaneously with both devices at the same time.

Measurements

Novel mHealth smartphone app. The mHealth app contained a purpose-built user interface (Figure 1) and was designed to monitor RR within the normal range. The user is instructed to hold their smartphone to their upper-middle chest with the screen facing outwards while sitting still and breathing normally for the duration of the 30-s sensor recording. Data is captured from the smartphone's tri-axial gyroscope and interpolated to achieve an even 100-Hz sample frequency. A low-pass Butterworth filter with 0.4 Hz cut-off is applied to remove high-frequency noise while retaining activity associated with breathing rates within the normal range typically in the 0.16-0.33-Hz range (10-20 breaths per minute (BPM)). RR is calculated by performing an autocorrelation before normalising the resulting signal. A peak-finding routine then identifies prominent peaks corresponding to the cyclical property of breathing movements. The mean inter-peak interval (IPI) is then calculated and converted to a 'per minute' RR estimation by division by 60 (seconds) (Figure 2).

An additional 'signal check' routine assesses whether the signal quality is sufficient to accurately derive RR, based on whether the number of autocorrelation peaks or standard deviation (*SD*) of the individual IPIs meets predetermined thresholds identified via preliminary bench-testing. If a recording fails the signal check, the user is informed via the app's UI, redirected to the operation instructions and prompted to try again. Passing the signal check within three recording attempts constitutes a successful use of the system, and three consecutive signal check failures constitute an unsuccessful use of the system, after which the user is instructed to seek support or try again later.

FDA-cleared reference

The *MightySat* Rx,¹⁴ developed by Masimo Corporation, was selected as a reference due to its FDA-cleared status, continuous measurement and ease of use. The fingertip pulse oximeter derives RR using photoplethysmography (PPG) (an optical measure of volumetric changes in peripheral blood flow). Continuous estimates of RR produced by this reference were converted to single-weighted averages to facilitate comparison with data derived from the mHealth app.



Figure 1. Selected wireframes from the user interface of the mHealth app, depicting (from left to right) an option given to the user to view operation instructions in video or written format, an instruction for the user to hold their smartphone to their chest, a clinical safety feature allowing the user to retake a recording if they were disturbed while taking the original recording, and feedback given to the user if their recording fails the signal check.



Figure 2. Graphical depiction of peak finding following the application of an autocorrelation method. The grey line shows an example of correlation coefficients for a movement sensor (gyroscope) signal correlated with itself at progressive temporal shifts. Black crosses indicate prominent peaks corresponding to the cyclical property of breathing movements. IPIs are depicted by *d_i*.

Novel reference

The novel reference method involves the identification of repeated cyclical peak-trough complexes within smartphone movement sensor signals (Figure 3). Signals of insufficient quality to derive RR are considered to fail the reference method. This method is conceptually similar to reference methods described in peer-reviewed literature



Figure 3. Graphical depiction of the novel reference method involving the visual inspection of smartphone movement sensor signals by trained clinical and scientific researchers. The grey solid line shows an example of a smartphone movement sensor (gyroscope) signal, with black solid arrows depicting nine full repeated cyclical peak-trough complexes. The grey dotted line indicates the projected continuation of the movement sensor signal past the end of the recording period, with the grey dashed arrow indicating where a tenth full peak-trough complex would end. Hence, the movement sensor signal depicts a total of approximately 9.75 peak-trough complexes, with the final .75 peak-trough complex indicated by the black dashed arrow.

reporting the accuracy assessments of multiple RR devices, including successful FDA market clearance applications.^{14,31–33} This method would permit accuracy assessments to be conducted via remote and real-world studies without a need for additional hardware, offering a significant value in terms of research scale, cost and ecological validity via the avoidance of observation bias.

Participants and recruitment

Participants were recruited via convenience sampling. All were employees of the mHealth app manufacturer. Inclusion criteria included were aged 18 or over and willing and able to follow the study protocol and complete an informed consent form.

Procedure

The study took place at the offices of the mHealth app manufacturer. Participants were provided with complete information concerning the study procedures and gave written informed consent to participate. The FDA-cleared reference device was applied to the forefinger of the participant's left hand. Participants were provided with an iPhone XR, model number MRY42B/A with the mHealth app installed and received verbal instructions on operating the device: namely, to hold the smartphone to their upper middle chest with the screen facing outwards while sitting still and breathing normally during the 30-s recording. Participants were instructed to capture six recordings, disregarding whether each recording passed or failed the signal check. Audiovisual footage was captured during the study and used for offline synchronisation of data captured via the mHealth app and FDA-cleared reference. Specifically, this included sounds produced by the mHealth app indicating the start and end of the app's recording period and depicting RR estimates displayed on the FDA-cleared reference's monitor. Participation took around 10 min per participant.

Statistics

The error of the mHealth app and novel reference relative to the FDA-cleared reference was assessed through measures of mean absolute error (MAE), relative MAE and using the Bland-Altman method.³⁴ Due to the non-normal distribution of absolute error data, confidence intervals (CIs) for MAE and relative MAE were derived via bootstrapping with replacement employing 1000 iterations and a sample size of 100%. The proportion of clinically significant errors, defined as an absolute error greater than three BPM,^{35,36} was also calculated. Direct relationships between RR estimates generated through the mHealth app, novel reference and FDA-cleared reference were assessed via Pearson Product Moment Correlation (PPMC). The usability of the mHealth app was assessed using the proportion and position of recordings that failed the signal check.

Results

Participants and data

Fifteen participants took part in Study 1 (nine females), for whom six recordings each were collected for a total of 90. Twenty-six (28%) mHealth app recordings failed the signal check and were excluded from analyses, resulting in a dataset of 64 paired samples. Twenty-nine (32%) of recordings failed the novel reference method, so were excluded from analyses, resulting in a dataset of 61 paired samples.

Accuracy

Mhealth app versus FDA-cleared reference. Error results indicated an MAE of 1.65 BPM (SD = 1.49) with a 95% (CI) of 1.32–2.06. Relative MAE was 12.2% (SD = 9.23) with 95% CI of 10.06–14.57. Bias (FDA-cleared reference–mHealth app) was 0.81 (SD = 2.08) with limits of agreement (LoA) of -3.27 to 4.89, indicating RR underestimation by the mHealth app. Eight comparisons (12.5%) had an absolute error greater than 3 BPM. A Bland–Altman plot indicated error values as a function of RR averaged between the reference and mHealth app (Figure 4). PPMC produced a coefficient of r(63) = 0.700, p < .000, indicating a high or strong association between the reference RR estimates and mHealth app RR estimate³⁷ (Figure 5).

Novel reference versus FDA-cleared reference

Error results indicated that MAE was 1.69 BPM (SD = 1.61) with a 95% CI of 1.23–2.22. Relative MAE was



Figure 4. Bland—Altman plot for RR estimates provided by the mHealth app and FDA-cleared reference. The *x*-axis indicates RR estimates averaged between the mHealth app and FDA-cleared reference, and the *y*-axis indicates the difference between RR estimates from each source (FDA-cleared reference—mHealth app). The solid horizontal line depicts a mean difference (bias) of 0, and dashed lines from top to bottom represent the upper limit of agreement (4.89), the observed mean difference (bias: 0.81), and the lower limit of agreement (—3.27). Marker size is proportional to the number of observations for each combination of values. FDA: Food and Drug Administration; RR: respiratory rate.



Figure 5. Scatterplot for simultaneous RR estimates provided by the mHealth app (*x*-axis) and FDA-cleared reference (*y*-axis). The solid line indicates the gradient y = x. Marker size is proportional to the number of observations for each combination of values. RR: respiratory rate.

12.8% (*SD* = 11.60) with 95% CI of 9.96–15.64. Bias (FDA-cleared reference–novel reference) was 0.22 (*SD* = 2.34) with LoA of -4.36 to 4.79, indicating slight RR underestimation by the mHealth app. Nine comparisons (15%) had an absolute error greater than 3 BPM. A Bland–Altman plot indicated error values as a function of RR averaged between the FDA-cleared and novel references (Figure 6). PPMC produced a coefficient of r(59) = 0.701, p < .000, indicating a high or strong association³⁷ (Figure 7).

Usability. Fourteen of 15 participants (93.3%) were able to use the system successfully on their first try (Table 1; Figure 8). Specifically, this indicates that they could capture one or more recordings that passed the signal check within the first three attempts. All participants were able to use the system successfully by the end of their second try.

Interim conclusion

Study 1 results indicated strong relationships between the FDA-cleared reference and both the mHealth app and the novel reference. Notably, these relationships were highly comparable to functional outcomes for alternative FDA-cleared RR monitoring devices.^{13,15,36,38} Accordingly, these results supported both the continued assessment of the mHealth app and the application of the novel reference to accuracy analyses in Study 2, as described below.

Study 2

Methods

Study 2 aimed to establish the accuracy of the mHealth app 'in the wild' via remote data capture, compared to the novel



Figure 6. Bland—Altman plot for RR estimates provided by the novel reference and FDA-cleared reference. The *x*-axis indicates RR estimates averaged between the novel and FDA-cleared references, and the *y*-axis indicates the difference between RR estimates from each source (FDA-cleared reference—novel reference). The solid horizontal line depicts a mean difference (bias) of 0, and dashed lines from top to bottom represent the upper limit of agreement (4.79), the observed mean difference (bias: 0.22), and the lower limit of agreement (-4.36). Marker size is proportional to the number of observations for each combination of values. FDA: Food and Drug Administration; RR: respiratory rate.

Table 1. Number and proportion of participants able to use the system successfully on consecutive attempts in Study 1 (n = 15).

	Recording attempt			
Use of system, n (%)	First	Second	Third	Total
First	11 (73.3)	12 (80.0)	10 (66.7)	14 (93.3)
Second	12 (80.0)	8 (53.3)	11 (73.3)	15 (100)

reference validated in the Study 1. The usability of the mHealth app was additionally assessed in a larger sample. Measures and statistics were as described for Study 1.

Participants and recruitment

Participants were recruited via an online research platform, with study enrolment controlled to ensure a proportionate distribution of age, gender and smartphone ownership (iOS vs Android). Inclusion criteria included were being aged 18 or over, having access to a smartphone of minimum requirements to download the mHealth app and being willing and able to follow the study protocol and complete an informed consent form. As researchers would not monitor participants during their participation, additional safety criteria excluded individuals who were pregnant and breastfeeding, had a pacemaker



Figure 7. Scatterplot for simultaneous RR estimates provided by the novel reference (*x*-axis) and FDA-cleared reference (*y*-axis). The solid line indicates the gradient y = x. Marker size is proportional to the number of observations for each combination of values. RR: respiratory rate.



Figure 8. Line graph indicating the proportion of Study 1 participants who were able to generate a recording that passed the signal check on each of six consecutive recording attempts. The black dashed line indicates the proportion of individuals who were able to generate a recording that passed the signal check by the end of their first use of the system (three consecutive recordings), and the grey dashed line indicates the proportion of individuals who were able to do so by the end of their second use of the system.

or self-reported a condition that might affect their breathing, such as asthma, or might affect their movement such as tremor.

Procedure

Participants were directed to online documentation containing full information about the study procedures before completing an online eConsent procedure. They then completed a baseline questionnaire concerning their demographics, including age, sex, ethnicity, height and weight, before receiving instructions to download and activate the mHealth app. Participants were requested to follow instructions provided within the mHealth app to capture 10 RR recordings, including recordings that both passed and failed the signal check, before completing a System Usability Scale (SUS)³⁹ and providing separate qualitative feedback on their experience using the mHealth app. Study-specific procedures took approximately 20 min, for which participants were reimbursed £2.50 through the research platform.

Results

Participants and data

One Hundred and sixty-five participants enrolled in the study, of whom 152 completed the baseline questionnaire concerning their demographics (Table 2). Medical conditions reported included asthma (respiratory), arthritis and Parkinson's disease (movement). Five participants were excluded due to significant deviation from the study protocol, resulting in a participant cohort of 160, for whom a mode of 11 mHealth app recordings each was captured. Nine hundred and eighty-seven recordings passed the signal check and were included in accuracy analyses. Recordings were submitted from 64 unique smartphone models, 46 (71.9%) of which were Android and the rest were iPhone models.

Accuracy

Error results indicated an MAE of 1.14 BPM (SD = 1.44) with a 95% CI of 1.02–1.26. Relative MAE was 9.5% (SD = 18.70) with 95% CI of 8.38–10.72. Bias (novel reference–mHealth app) was 0.08 (SD = 1.84) with LoA of –3.68 to 3.51, indicating slight RR underestimation by the mHealth app. Sixty-one comparisons (6.2%) had an absolute error greater than 3 BPM. No difference in MAE was found between Android and Apple devices. A Bland–Altman plot indicated error values as a function of RR averaged between the reference and mHealth app (Figure 9).

PPMC produced a coefficient of r(986) = 0.855, p < .000, indicating a high or strong association³⁷ (Figure 10).

Usability

One Hundred and forty-nine (93.1%) of a total of 160 participants who captured mHealth app recordings were able to use the system successfully on their first try (Table 3; Figure 11). One Hundred and fifty-five (96.9%) did so by their second try.

The mean SUS score was 73.2 (SD = 5.39). Of the subscales, each scored between 0 and 4, those most agreed with by participants were: I would imagine that most people would learn to use this system very quickly (3.2), I thought the system was easy to use (3.1) and I felt very confident using the system (3.0). The lowest scoring, indicating participant disagreement, was: I thought that I would need

Table 2. Demographic characteristics f	or Study 2 participants (<i>n</i> = 152)
--	---

Characteristic	Female ($n = 67$)	Male ($n = 85$)	Total (<i>n</i> = 152)
Age in years, mean (SD, range)	43.5 (14.85, 18–73)	40.7 (14.70, 19-69)	41.9 (14.78, 18–73)
Weight in kg, mean (SD, range)	70.2 (19.45, 47–159)	87.9 (22.31, 52–203)	80.1 (22.08, 47–203)
Height in m, mean (SD, range)	1.66 (0.108, 1.45-2.16)	1.78 (0.660, 1.56-1.93)	1.73 (1.061, 1.45–2.16)
BMI in kg/m², mean (SD, range)	25.4 (6.28, 17.8-58.4)	27.7 (6.80, 17.4–62.5)	26.7 (6.65, 17.4-62.7)
Ethnicity, n (%)			
White	57 (85.1)	72 (84.7)	129 (84.9)
Asian	4 (6.0)	7 (8.2)	11 (7.2)
Black	3 (4.5)	2 (2.4)	5 (3.3)
Mixed/multiple	3 (4.5)	4 (4.7)	7 (4.6)
Medical conditions, n (%)			
Respiratory disorder	9 (13.4)	7 (8.2)	16 (10.5)
Movement disorder	2 (3.0)	0 (0)	2 (1.3)
Cognitive disorder	0 (0)	0 (0)	0 (0)



Figure 9. Bland—Altman plot for RR estimates provided by the mHealth app and novel reference. The *x*-axis indicates RR estimates averaged between the mHealth app and novel reference, and the *y*-axis indicates the difference between RR estimates from each source (novel reference—mHealth app). The solid horizontal line depicts a mean difference (bias) of 0, and dashed lines from top to bottom represent the upper limit of agreement (3.51), the observed mean difference (bias: 0.08), and the lower limit of agreement (-3.68). Marker size is proportional to the number of observations for each combination of values. RR: respiratory rate.

the support of a technical person to be able to use this system (0.5) and I needed to learn a lot of things before I could get going with this system (0.8).



Figure 10. Scatterplot for simultaneous RR estimates provided by the mHealth app (*x*-axis) and novel reference (*y*-axis). The solid line indicates the gradient y = x. Marker size is proportional to the number of observations for each combination of values. RR: respiratory rate.

Discussion

Principal findings

To the authors' knowledge, this is the first technical validation to assess at scale a user-operated novel mHealth smartphone application designed to capture a user's RR using smartphone movement sensors, considering both accuracy and usability in

Table 3. Number and proportion of participants able to use the system successfully on consecutive attempts in Study 2 (n = 160).

	Recording attempt			
Use of system, n (%)	First	Second	Third	Total
First	102 (63.8)	117 (74.1)	119 (77.3)	149 (93.1)
Second	102 (66.2)	104 (68.4)	105 (70.0)	155 (96.9)



Figure 11. Line graph indicating the proportion of Study 2 participants who were able to generate a recording that passed the signal check on each of ten consecutive recording attempts. The black dashed line indicates the proportion of individuals who were able to generate a recording that passed the signal check by the end of their first use of the system (three consecutive recordings), and the grey dashed line indicates the proportion of individuals who were able to do so by the end of their second use of the system.

an ecologically valid study environment. Outcomes for the mHealth app were highly comparable to results published for medical devices available on the market today (Table 4). In addition, as changes in breathing rate greater than 3 BPM may indicate clinical deterioration,^{35,36} observations that error values for the mHealth app were typically less than this threshold suggest the device may carry low clinical risk. Study 2 revealed a small cluster of substantial overestimation errors (5-10 BPM) for lower RRs (8-14 BPM). Although this observation was not found in Study 1, this may be due to the smaller sample size in that analysis. The nature of these overestimations is unclear based on the present analyses. The overestimation of RR carries clinical risk with regard to both the underdiagnosis of bradypnea (low RR) and the overdiagnosis of tachypnea (elevated RR) that may lead to clinical decisionmaking based on misinformation, although it should be noted that RR is rarely used in isolation to inform clinical decisionmaking. Future research should seek to identify and mitigate the cause of these errors.

Concerning usability, most participants could successfully operate the mHealth app on their first or second use of the system. Although no industry standards for successful operation exist, results observed here appeared to be broadly similar to values that could be estimated from the available literature regarding other physiological measurement mHealth apps, which were typically in the range of 95% or higher. 40-42 Subjective usability outcomes were also promising, with an overall SUS score well above the industry average of 68.43 Study 2 revealed a general trend of high signal check pass rates for later sequential recording attempts, suggesting that participants found it easier to capture RR recordings the more they used the mHealth app. Although this learning effect was not observed in the Study 1 results, this may be due to observer bias and a small sample size within that study setting. This observation holds promise for improved usability with the longterm use of the mHealth app, although it may indicate greater clinical risk during the early use of the system. Future research may seek to steepen the learning curve to minimise clinical risk.

Wearable devices equipped with PPG sensors provide another alternative method for low cost and the remote measurement of RR. However, PPG methods have hardware and processing power requirements that are higher than smartphone movement sensor methods. Smartphone devices are also used more widely and therefore have greater availability in remote locations or low income countries.⁴⁴

Strengths of the present study include the application of a remote study design that lends ecological validity to the results and selective recruitment to ensure a heterogeneous participant cohort, which suggests good generalisability of the results. In addition, the inclusion of usability assessment allows a holistic perspective on the mHealth app to be generated. In all, these results hold promise for the use of smartphone movement sensors as a viable means of remote RR monitoring. Software-based mHealth may offer cost and scalability benefits compared to hardware-based monitoring. Additionally, movement sensors may better protect RR signal quality than alternative devices that use microphone and camera sensors, as these are vulnerable to noise from environmental light and sound that is difficult to control. These benefits suggest that RR monitoring based on smartphone movement sensors may support healthcare systems to care for their patients when they are outside of the clinic better than currently available alternatives.

Study limitations

As all participants had RR within the normal range, it is unclear how the observed results may extrapolate to healthcare patients who would be likely real-world users of the mHealth app, particularly those with abnormal breathing rates and patterns due to a respiratory condition. For example, individuals receiving ventolin for asthma may

Comparison	MAE (<i>SD</i> , 95% CI)	Relative MAE (<i>SD</i> , 95% Cl)	Bias (<i>SD</i> , LoA)	Correlation coefficient
mHealth app				
Compared to FDA-cleared reference	1.65 (1.49, 1.32 —2.06)	12.2 (9.23, 10.06 —14.57)	0.81 (2.08, -3.27-4.89)	0.700
Compared to novel reference	1.14 (1.44, 1.02 —1.26)	9.5 (18.70, 8.38 —10.72)	0.08 (1.84, -3.68-3.51)	0.885
Respirasense (PDM Solutions)				
Compared to capnography ¹³	-	-	0.38 (N/A, N/A, -1.0 to 1.8)	-
Compared to manual count ¹³	-	-	-0.70 (N/A, N/A, -4.9 to 3.5)	-
Compared to electrocardiogram ³⁶	-	-	-0.41 (1.79, -0.73 to -0.08, -3.9 to 3.1)	0.84
Compared to manual count ³⁶	-	-	-0.58 (2.5, -1.04 to -0.12, -5.5 to 4.3)	0.78
BioStamp nPoint (MC10) compared to capnography ¹⁵	1.3 (2.1, N/A)	-	-0.29 (N/A, N/A, -5.17 to 4.59)	0.697
Rad-87 (Masimo) compared to capnography ³⁷	-	10 (9, 7-13)	-	-

Table 4. Comparison of mHealth app results to alternative devices available on the market today.

have a medication induced tremor, which may affect gyroscope recordings. This is also true for individuals with disorders that are associated with tremor such as Parkinson's disease. Additionally, measuring RR using the mHealth app in the presence of a chronic, persistent cough, like those associated with severe asthma or COPD may require additional signal processing considerations. Additional signal processing considerations may be required for clinical use cases that require the detection of tachypnea or bradypnea. Participants both from Study 1 (employees of the mHealth app manufacturer) and Study 2 (members of an online research community) were likely to be technologically confident and may have therefore been predisposed to successfully operating the mHealth app. Future research should seek to incorporate individuals of low technical literacy and target end-users with relevant medical conditions to better understand these results' generalisability.

Concerning methodology, the FDA-cleared reference used in Study 1 has its own measurement error.¹⁴ Hence, error estimates presented here are, in fact, an unknown combination of errors associated with the FDA-cleared reference and mHealth app versus true RR. The Study 2 reference also underwent only limited validation in Study 1 and should be assessed more rigorously. Future research may wish to apply a wider range of reference methods, including gold and industry-standard references, to reduce the vulnerability of the mHealth app to shortcomings of any single reference.

Additionally, the present research design does not directly address potential benefits the mHealth app may offer if applied in a healthcare setting. Although expectations that moving health assessments outside of a clinical setting via mHealth technologies will improve healthcare economics have been somewhat supported by literature,⁵ clinical evidence suggests that mHealth technologies are highly heterogeneous in their ability to improve health outcomes.^{45,46} Suggestions that mHealth may help to overcome social, economic and geographical barriers to healthcare are also yet to be validated.^{47–49} Future research should seek to understand the clinical, economic and social outcomes associated with real-world use of the mHealth app.

Conclusions

Decentralised healthcare technology holds the potential to offer clinical and economic benefits to patients, HCPs and healthcare systems. Breathing is an important indicator of health, and although solutions for remote RR monitoring exist, many entail significant shortcomings that limit their ability to capitalise on potential benefits of mHealth. Results from this technical validation hold promise for the use of smartphone movement sensors as a robust means for remote RR monitoring. However, future research should address residual questions and risks associated with the technology identified in this article and seek to validate the impact of similar technologies as applied in the real world.

Acknowledgements: This research was sponsored by Huma Therapeutics Ltd. The authors are grateful to Huma's wonderful developers Michele Colombo, Stanislas Heili, Leonardo Festa, Davide Mascitti, Emanuele Distefano, Matteo Puccinelli and Matteo Vigoni for their diligent efforts in preparing the mHealth app for this research. Additional thanks to Emily Binning for her continued strategic support that helped this research come to fruition.

Declaration of conflicting interests: The author(s) declared the following potential conflicts of interest with respect to the research, authorship and/or publication of this article. All authors are the current or previous employees of Huma Therapeutics Ltd, which is the developer of the mHealth smartphone app. No additional conflicts of interest relevant to this study are declared.

Funding: The author(s) disclosed receipt of the following financial support for the research, authorship and/or publication of this article: This work was supported by Huma Therapeutics Ltd (Grant No. N/A).

Author contributions: SV, AC, BK, MD and MB drafted the manuscript, conducted statistical analyses and interpreted the results. DP aided in statistical analysis, interpreted the results and revised the manuscript for intellectual content. MA and DP revised the manuscript for intellectual content. All authors approved the final manuscript for submission.

Guarantor: All authors are willing to take full responsibility for the article, including for the accuracy and appropriateness of the reference list.

Ethical approval: Ethical approval was provided by the University of Exeter's Research Ethics Board (application ID: eUEBS004088), and all research was conducted in compliance with the Declaration of Helsinki.

Informed consent: Not applicable, because this article does not contain any studies with human or animal subjects.

Trial registration: Not applicable, because this article does not contain any clinical trials.

ORCID iD: David Plans D https://orcid.org/0000-0002-0476-3342

References

1. Duggal R, Brindle I and Bagenal J. Digital healthcare: regulating the revolution. *Br Med J* 2018; 360: k6.

- Morgan AU, Balachandran M and Do D. Remote monitoring of patients with COVID-19: design, implementation, and outcomes of the first 3,000 patients in COVID watch. *NEJM Catal Innov Care Deliv* 2020; 1. DOI: 10.1056/CAT.20. 0342. Epub ahead of print.
- Watson AR, Wah R and Thamman R. The value of remote monitoring for the COVID-19 pandemic. *Telemed J E Health* 2020; 26: 1110–1112.
- 4. Shah SS, Gvozdanovic A, Knight M, et al. Mobile app-based remote patient monitoring in acute medical conditions: prospective feasibility study exploring digital health solutions on clinical workload during the COVID crisis. *JMIR Form Res* 2021; 5: e23190.
- Iribarren SJ, Cato K, Falzon L, et al. What is the economic evidence for mHealth? A systematic review of economic evaluations of mHealth solutions. *PLoS One* 2017; 12. DOI: 10. 1371/journal.pone.0170581. Epub ahead of print.
- Marcolino MS, Oliveira JAQ, D'Agostino M, et al. The impact of mHealth interventions: systematic review of systematic reviews. *JMIR Mhealth Uhealth* 2018; 6. DOI: 10. 2196/mhealth.8873. Epub ahead of print.
- Gravelyn TR and Weg JG. Respiratory rate as an indicator of acute respiratory dysfunction. JAMA 1980; 244: 1123–1125.
- Chen L, Reisner AT, Gribok A, et al. Can we improve the clinical utility of respiratory rate as a monitored vital sign? *Shock* 2009; 31: 574–580.
- Fieselmann JF, Hendryx MS, Helms CM, et al. Respiratory rate predicts cardiopulmonary arrest for internal medicine inpatients. J Gen Intern Med 1993; 8: 354–360.
- Miller DJ, Capodilupo JV and Lastella M. Analyzing changes in respiratory rate to predict the risk of COVID-19 infection. *PLoS One* 2020; 15. DOI: 10.1371/journal.pone.0243693. Epub ahead of print 2020.
- 11. Roca O, Caralt B and Messika J. An Index combining respiratory rate and oxygenation to predict outcome of nasal high-flow therapy. *Am J Respir Crit Care Med* 2019; 199: 1368–1376.
- Myint PK, Musonda P and Sankaran P. Confusion, urea, respiratory rate and shock Index or adjusted shock Index (CURSI or CURASI) criteria predict mortality in community-acquired pneumonia. *Eur J Intern Med* 2010; 21: 429–433.
- Subbe CP and Kinsella S. Continuous monitoring of respiratory rate in emergency admissions: evaluation of the RespiraSenseTM sensor in acute care compared to the industry standard and gold standard. *Sensors* 2018; 18. DOI: 10. 3390/s18082700. Epub ahead of print.
- U.S. Food and Drug Administration. Response to 510(k) Premarket Notification for Masimo MightySat Rx Fingertip Pulse Oximeter (K181956). https://www.accessdata.fda.gov/ cdrh_docs/pdf18/K181956.pdf (2018, accessed 1 June 2021).
- Sen-Gupta E, Wright DE and Caccese JW. A pivotal study to validate the performance of a novel wearable sensor and system for biometric monitoring in clinical and remote environments. *Digit Biomark* 2019; 3: 1–13.
- Liu H, Allen J, Zheng D, et al. Recent development of respiratory rate measurement technologies. *Physiol Meas* 2019; 40. DOI: 10.1088/1361-6579/ab299e. Epub ahead of print.
- Khalid A. Pulse oximeters are selling out because of the pandemic. Most people don't need them. https://qz.com/1832464/

pulse-oximeters-for-coronavirusunnecessary-but-selling-strong/ (2020, accessed 1 June 2021).

- National Advisory Group on Health Information Technology in England. Making IT work: harnessing the power of health information technology to improve care in England. https://assets.publishing.service.gov.uk/government/uploads/ system/uploads/attachment_data/file/550866/Wachter_Review_ Accessible.pdf (2016, accessed 1 June 2021).
- Alafeef M and Fraiwan M. Smartphone-based respiratory rate estimation using photoplethysmographic imaging and discrete wavelet transform. *J Ambient Intell Humaniz Comput* 2020; 11: 693–703.
- Nam Y, Lee J and Chon KH. Respiratory rate estimation from the built-in cameras of smartphones and tablets. *Ann Biomed Eng* 2014; 42: 885–898.
- Sanyal S and Nundy KK. Algorithms for monitoring heart rate and respiratory rate from the video of a user's face. *IEEE J Transl Eng Health Med* 2018; 6: 1–11. DOI: 10.1109/ JTEHM.2018.2818687.
- Phokela KK and Naik V. Use of smartphone's headset microphone to estimate the rate of respiration. In: 2020 International Conference on COMmunication Systems & NETworkS (COMSNETS), 2020, pp. 64–69. DOI: 10.1109/ COMSNETS48256.2020.9027297.
- Jin A, Yin B, Morren G, et al. Performance evaluation of a triaxial accelerometry-based respiration monitoring for ambient assisted living. *Annu Int Conf IEEE Eng Med Biol Soc* 2009; 2009: 5677–5680. DOI: 10.1109/IEMBS.2009.5333116.
- Nurmi S, Saaresranta T, Koivisto T, et al. Validation of an accelerometer based BCG method for sleep analysis. Espoo: Aalto University, 2016. ISBN:9789526068428
- Jarchi D, Rodgers SJ, Tarassenko L, et al. Accelerometrybased estimation of respiratory rate for post-intensive care patient monitoring. *IEEE Sens J* 2018; 18: 4981–4989.
- Hernandez J, Li Y, Rehg JM, et al. BioGlass: physiological parameter estimation using a head-mounted wearable device. In: 2014 4th International Conference on Wireless Mobile Communication and Healthcare Transforming Healthcare Through Innovations in Mobile and Wireless Technologies (MOBIHEALTH), 2014, pp. 55–58. DOI: 10.1109/MOBIHEALTH.2014.7015908.
- Shen C-L, Huang T-H and Hsu P-C. Respiratory rate estimation by using ECG, impedance, and motion sensing in smart clothing. *J Med Biol Eng* 2017; 37: 826–842.
- Aly H and Youssef M. Zephyr: Ubiquitous accurate multisensor fusion-based respiratory rate estimation using smartphones. In: *IEEE INFOCOM 2016 - The 35th Annual IEEE International Conference on Computer Communications*, 2016, pp. 1–9. DOI: 10.1109/INFOCOM.2016.7524401.
- Rahman MM, Nemati E, Nathan V, et al. Instantaneous respiratory rate estimation on context-aware mobile devices. In: 13th EAI International Conference on Body Area Networks. Springer International Publishing, pp. 267–281.
- Li X. Using mobile phone sensors to detect rapid respiratory rate in the diagnosis of pneumonia. *IACSIT Int J Eng Technol* 2016; 8: 293–296.
- U.S. Food and Drug Administration. Response to 510(k) Premarket Notification for Masimo Centroid System (K191882). https://www.accessdata.fda.gov/cdrh_docs/pdf19/ K191882.pdf (2020, accessed 1 June 2021).

- Tal A, Shinar Z, Shaki D, et al. Validation of contact-free sleep monitoring device with comparison to polysomnography. *J Clin Sleep Med* 2017; 13: 517–522.
- U.S. Food and Drug Administration. Response to 510(k) Premarket Notification for EarlySense Bed Sensing Unit (K171836). https://www.accessdata.fda.gov/cdrh_docs/pdf17/ K171836.pdf (2017, accessed 1 June 2021).
- Bland JM and Altman DG. Measuring agreement in method comparison studies. *Stat Methods Med Res* 1999; 8: 135–160.
- 35. Dougherty L and Lister S. *The royal marsden manual of clinical nursing procedures*. John Wiley & Sons, 2015.
- Lee PJ. Clinical evaluation of a novel respiratory rate monitor. J Clin Monit Comput 2016; 30: 175–183.
- Mason R, Lind D and Marchal W. *Statistics: an introduction*. Wisconsin: Cole Publishing Company, 1998.
- Ramsay MAE, Usman M, Lagow E, et al. The accuracy, precision and reliability of measuring ventilatory rate and detecting ventilatory pause by rainbow acoustic monitoring and capnometry. *Anesth Analg* 2013; 117: 69–75.
- 39. Brooke J. SUS: a "quick and dirty" usability scale. Philadelphia, PA: CRC Press, 1996. ISBN:9780748404605
- 40. Yan BP, Chan CK and Li CK. Resting and postexercise heart rate detection from fingertip and facial photoplethysmography using a smartphone camera: a validation study. *JMIR Mhealth Uhealth* 2017; 5. DOI: 10.2196/mhealth.7275. Epub ahead of print.
- Coppetti T, Brauchlin A and Müggler S. Accuracy of smartphone apps for heart rate measurement. *Eur J Prev Cardiol* 2017; 24: 1287–1293.
- Parpinel M, Scherling L, Lazzer S, et al. Reliability of heart rate mobile apps in young healthy adults: exploratory study and research directions. *BMJ Health Care Inform* 2017; 24. DOI: 10.14236/jhi.v24i2.921.
- Lewis SJRJ. Item benchmarks for the system usability scale. J Usability Stud 2018; 13: 158–167.
- Natarajan A, Su H-W, Heneghan C, et al. Measurement of respiratory rate using wearable devices and applications to COVID-19 detection. *Npj Digit Med* 2021; 4: 1–10.
- 45. Free C, Phillips G, Galli L, et al. The effectiveness of mobilehealth technology-based health behaviour change or disease management interventions for health care consumers: a systematic review. *PLoS Med* 2013; 10(1): e1001362. DOI: 10. 1371/journal.pmed.1001362.
- Carreiro S, Newcomb M, Leach R, et al. Current reporting of usability and impact of mHealth interventions for substance use disorder: a systematic review. *Drug Alcohol Depend* 2020; 215: 108201. DOI: 10.1016/j.drugalcdep. 2020.108201.
- 47. Hong AY and Cho J. Has the digital health divide widened? Trends of health-related internet use among older adults from 2003 to 2011. *J Gerontol B Psychol Sci Soc Sci* 2016; 72: 856–863.
- Hall CS, Fottrell E, Wilkinson S, et al. Assessing the impact of mHealth interventions in low- and middle-income countries– what has been shown to work? *Glob Health Action* 2014; 7: 25606. DOI: 10.3402/gha.v7.25606.
- Crawford A and Serhal E. Digital health equity and COVID-19: the innovation curve cannot reinforce the social gradient of health. *J Med Internet Res* 2020; 22(6): e19361. DOI: 10.2196/19361.