

1 Mini-Review

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3 **‘Smart Diagnosis’ of Parasites using Smartphones**

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11 **ABSTRACT** Accurate and rapid diagnosis is crucial in combating parasitic diseases that  
12 cause millions of deaths worldwide. However, the scarcity of specialized diagnostic  
13 equipment in low-to-middle income countries is one of the barriers in effective management  
14 of parasitic diseases, which warrants the need for alternative, inexpensive and point-of-care  
15 diagnostic tools. Due to their multiple built-in sensors, smartphones offer cost-effective  
16 alternative to expensive diagnostic devices. However, the use of smartphones in parasitic  
17 diagnoses remains in its infancy. This mini-review describes various smartphone-based  
18 devices, applied specifically for the diagnosis of parasitic diseases and discusses challenges  
19 and potential implications for their use in future.

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22 **KEYWORDS** Smartphones, parasitic diseases, diagnosis, microscopy, lab-on-a-chip

23 Parasitic diseases cause millions of morbidities and mortalities, and impose serious health  
24 and socioeconomic consequences, mainly in developing countries of the world (1).  
25 According to the Centers for Disease Control and Prevention, malaria alone causes  
26 approximately 660,000 casualties per annum, and the Neglected Tropical Diseases (NTDs),  
27 including Chagas disease, echinococcosis, schistosomiasis, soil-transmitted helminthiases,  
28 African trypanosomiasis, cysticercosis, lymphatic filariasis, scabies etc. affect millions of  
29 people worldwide (2). Accurate and rapid diagnosis is of paramount importance in the  
30 effective clinical management of such parasitic diseases. However, the diagnosis of parasitic  
31 diseases is severely compromised due to the scarcity of trained personnel, and the lack of  
32 specialized diagnostic equipment in developing countries. For instance, the utility of many  
33 commonly used diagnostic methods for parasitic diseases such as microscopy and nucleic  
34 acid amplification is hindered by the unavailability of the skilled workforce, expensive  
35 instrument(s), reagents and electricity in developing countries (1, 3). This situation results in  
36 the inadequacy of these diagnostic tools for the neediest communities, leading to a  
37 compromise in the management of parasitic diseases.

38 Mobile phones and smartphones have brought enormous convenience and sizable impact  
39 to the modern society as depicted by a wide range of smartphone users worldwide.  
40 Smartphones are the more advanced form of mobile phones with fully functional computing  
41 capabilities and user-friendly features such as personal information management applications,  
42 compact digital cameras, Global Positioning System (GPS) navigation, internet access etc.  
43 Most smartphones are designed to have multiple sensors such as imaging camera, vibration  
44 sensor, GPS sensor and light level sensor etc. (4). Due to these powerful built-in-sensors,  
45 smartphones are setting their roots in the medical field as an alternative to expensive  
46 laboratory instruments for various diagnostic purposes (4), and they are of particular interest  
47 in regions with limited resources (5). However, the use of smartphones and mobile devices in

48 the diagnosis of parasitic diseases remains in its infancy and limited information is available  
49 on the topic. This article is aimed to review all available studies (on 16-06-2017) on hardware  
50 or software components of smartphones, applied specifically to address the diagnosis of  
51 parasites of medical or veterinary importance. Furthermore, the review discusses future  
52 implications and challenges with reference to parasitic diseases.

53

## 54 **LITERATURE SEARCH**

55 Using ISI Web of Knowledge, all the databases were searched from 1900 to 2017  
56 (accessed on June 16, 2017) with multiple search terms and filters (see Supplementary Table  
57 1). The same terms were used to search articles on PubMed and Google Scholar. Additional  
58 relevant articles were identified from the references cited in the articles found in the primary  
59 search. Twenty-four studies (Table 1) related to the smartphone-based diagnosis of parasites  
60 were finally included in this review.

61

## 62 **PARASITE DIAGNOSIS USING SMARTPHONES**

63 This section provides the design and applications of various smartphone-based diagnostic  
64 methods and devices used for the diagnoses of parasites. For the convenience of readers, we  
65 have created various categories for different devices; however, some of them may fall under  
66 more than one categories (see Tables 1 and 2).

67 **Smartphone standalone technology.** Owing to high magnification lenses and powerful  
68 image processors, a smartphone standalone (i.e., without using any external enhancement  
69 such as a lens or a microscope) presents a useful tool for the diagnosis of parasitic diseases.  
70 For example, Meena and Bhatia used the smartphone for the first time to diagnose a cestode  
71 parasite in tomographic images (see Table 1), and they examined images of a small

72 cysticercus (a larval stage of a cestode) using the smartphone which was otherwise invisible  
73 to the clinicians on visual examination (6).

74 Smartphone applications (apps) and algorithms present another use of smartphones as a  
75 standalone tool in the diagnosis of parasitic diseases such as the interpretation of the rapid  
76 diagnostic test (RDT) results for malaria (7, 8). Although RDTs present an inexpensive point-  
77 of-care (POC) tool, their effective application in the diagnosis of malarial parasites could be  
78 impeded by an incorrect analysis of the results by a poorly-trained end-user (7). To avoid  
79 visual interpretation, a smartphone was used to image and transfer the results of the RDTs of  
80 malaria to the REDCap (a globally accessible database) for analysis using a specialized  
81 algorithm. Despite its slightly lower sensitivity (see Table 1), this method significantly  
82 reduced reporting errors and false-negative diagnosis compared to a method of visual  
83 interpretation (7). In another study, the control line on a malaria RDT was converted into a  
84 smartphone-readable quick response (QR) code (8). A smartphone was deployed to capture  
85 RDT images, and an associated app was used to perform image processing and recognition of  
86 QR code to determine the concentration of histidine-rich protein 2 (a *Plasmodium falciparum*  
87 specific protein). The detection limit of the assay was 0.966 nM (~543 parasites per  $\mu\text{L}$ )  
88 compared to that of the World Health Organization (WHO) benchmark testing for an RDT  
89 (500 parasites per  $\mu\text{L}$ ) for low parasitaemia, suggesting that this method needed modification  
90 to increase its sensitivity. Overall, these smartphone-based diagnostic techniques allow  
91 automated identification, secured record keeping, and quality assurance that could be highly  
92 useful in malaria surveillance programs.

93 Although thousands of smartphone apps are currently being used in the healthcare  
94 industry, there is a limited information available on apps for the diagnosis of parasites.  
95 TickID (<https://itunes.apple.com/US/app/id531348104>) is an example of such a free-  
96 downloadable app for smartphones (9). This app provides basic information on identification

97 (pictures of male, female and juvenile ticks) and management (disease biology, personal  
98 protection, tick-removal etc.) of selected ticks and tick-borne diseases for a common user.  
99 Similar smartphone apps could be developed for socioeconomically important parasites  
100 regionally as well as globally that could present a great assistance for the diagnosis and  
101 management of parasitic diseases.

102 **Lens-mounted smartphone ‘microscopy’.** Mounting a simple, portable lens on a  
103 smartphone camera can provide a powerful handheld microscope for the identification of  
104 parasites. The lens size determines the spatial resolution and field of view (FOV) as smaller  
105 lenses have smaller FOV but the greater spatial resolution and vice versa (29). Bogoch et al.  
106 constructed a handheld microscope by mounting a 3 mm ball-lens to a smartphone camera,  
107 and used it for the identification of soil-transmitted helminth (STH) and *Schistosoma* eggs in  
108 urine and stool samples of school-aged children (10, 11). Although this device showed low-  
109 to-moderate sensitivities and specificities, (see Table 1), and had small FOV producing  
110 inferior quality images (see Table 2), this is an inexpensive and portable microscope. With  
111 improved sensitivity, this could be invaluable in the field diagnosis of STH in developing  
112 countries.

113 In order to increase the resolution of lens-mounted smartphone microscopy, Switz et al.  
114 applied a reversed camera lens to a smartphone to produce a large FOV ( $\sim 10 \text{ mm}^2$ ) with a  
115 resolution ( $\leq 5 \mu\text{m}$ ) for better quality images of STH eggs in stool samples (12). A major issue  
116 in imaging parasitic eggs is their scattering at different focal depths in a three dimensional  
117 (3D) plane. Sowerby et al. addressed this issue by mounting a 12 mm double-convex  
118 objective lens on a smartphone camera to image *Ascaris lumbricoides* eggs and created  
119 composite images, using a software, ImageJ (13). Overall, external lens-mounted smartphone  
120 microscopes are portable, inexpensive, and operate without constant electricity needs, which

121 make them a field-deployable tool in parasitic diagnosis in resource-constrained regions of  
122 the world.

123 **Smartphone-assisted manual microscopy.** Smartphones have recently been applied in  
124 conjunction with various microscopic assemblies for the diagnosis of parasites. Ephraim et al.  
125 used smartphone-assisted Foldscope and reversed-lens CellScope for the diagnosis of  
126 *Schistosoma haematobium* eggs in urine samples of school-aged children (14). The handheld  
127 Foldscope was paper-made, consisted a 2.38 mm ball-lens, and a light-emitting diode (LED)  
128 secured to a smartphone camera. The reversed-lens CellScope was constructed with a lens  
129 embedded in a 3D-printed plastic and secured to a smartphone camera. Despite their low-to-  
130 moderate sensitivities (see Table 1), both ‘microscopes’ showed high specificities. In another  
131 field study, the CellScope consistently demonstrated high specificity despite low sensitivity  
132 for the diagnosis of *Schistosoma* eggs in urine and stool samples (15), indicating that with an  
133 enhanced sensitivity, these devices could be deployed in the field at a large-scale screening of  
134 schistosomes.

135 In an attempt to design a compact microscope, Tseng et al. introduced a lens-free  
136 microscope for the identification of *Giardia lamblia* cysts (16). The sample of interest was  
137 illuminated, using an incoherent LED light (shined vertically). The scattered light interfered  
138 with unscattered LED light to create a hologram of each cell, which was detected by a  
139 smartphone camera. Depending on the power of the smartphone, extremely rich information  
140 in hologram allows rapid reconstruction of the microscopic images. In another study, *G.*  
141 *lamblia* cysts were identified using a smartphone-based fluorescent microscope (17), where  
142 an LED light was used to excite the sample, and the emitted fluorescent light was detected  
143 with an external lens placed in front of a smartphone camera. For fluorescent imaging in this  
144 study, a dark-field background was created using an inexpensive color filter (17).

145 Smartphone-assisted microscopes have also been applied for the diagnosis of *Plasmodium*  
146 spp. For instance, a bright-field microscope was constructed using objective and wide-field  
147 eyepiece lenses to produce a magnification (28X) onto a smartphone camera for the  
148 identification of *P. falciparum* in blood smears (18). In another field study, *P. falciparum* was  
149 identified in Giemsa-stained blood films, using a handheld Newton Nm1 light microscope  
150 attached to a smartphone (19). The system achieved a moderate sensitivity and a high  
151 specificity (see Table 1), suggesting that this could be invaluable in large-scale malaria  
152 screening programs. Other malarial biomarkers such as hemozoin have also been detected in  
153 blood smears, using a low-cost and high-fidelity smartphone-assisted polarized microscope  
154 (20). However, this system requires adequate lens resolution to differentiate the presence of  
155 hemozoin within an infected blood smear.

156 **Smartphone-assisted automated microscopy.** Manual microscopic examination of  
157 parasitic eggs is considered laborious and time-consuming as it requires a microscope as well  
158 as a trained person, which limits its use in the field in developing countries. A possible  
159 solution to this problem could be the use of a dedicated smartphone app or algorithm for an  
160 automated detection of parasites. For instance, Linder et al. introduced two pattern  
161 recognition algorithms for the identification of *S. haematobium* eggs in images acquired by a  
162 smartphone or a webcam (21). This method achieved a high specificity and a moderate  
163 sensitivity, compared to visual identification method (see Table 1). In another study,  
164 Slusarewicz et al. introduced a smartphone-based fecal egg counting technique for animal  
165 parasites (22). The eggs were stained with a fluorescent chitin-binding protein and  
166 photographed using a smartphone, followed by automated egg counting with ImageJ. For  
167 strongyle eggs, a significant linear correlation ( $R^2 = 0.98$ ) and coefficient of variation were  
168 found between the automated counts and manual McMaster counts, indicating that the



169 automated system performs better than the most commonly used traditional method in  
170 veterinary parasitology.

171 Smartphone-assisted automated microscopy is not confined to egg identification only as a  
172 smartphone-based fluorescent microscopy has recently been applied to quantify DNA from  
173 *Trypanosoma cruzi* (25). PCR was performed inside a central processing unit (CPU) by  
174 controlling the heating/cooling cycles with a computer software. PCR products were exposed  
175 to UV light and imaged by a smartphone, using a low-cost filter. A histogram of the pixel  
176 intensities of the patient sample was compared to a control sample for detecting target  
177 pathogenic DNA (25). Similarly, Koydemir et al. designed a smartphone-fluorescent  
178 microscope with a large FOV ( $\sim 0.8 \text{ cm}^2$ ) to detect *G. lamblia* cysts (23). In this method, a  
179 smartphone was used to image fluorescently-labelled cysts, captured on a membrane, and the  
180 images were transferred to a remote processing system for automatic detection and counting  
181 of cysts with an algorithm in large volumes of water (e.g. 10-20 mL) in a short time (23).  
182 Rosado et al. have recently introduced a smartphone-based image processing and analysis  
183 methodology for identification of *P. falciparum* trophozoites in Giemsa-stained blood smears  
184 (24). The system automatically identified the parasite based on pre-annotated characters and  
185 achieved a moderate sensitivity and a high specificity (see Table 1).

186 Smartphone-assisted microscopy is not confined to still-imaging only as the use of  
187 smartphone video microscopy has been demonstrated recently for the quantification of *Loa*  
188 *loa* microfilariae (a larval stage of the parasite that has a serpentine movement) (26). The  
189 device (CellScope Loa) used a smartphone to perform video imaging of an unprocessed  
190 blood sample which was analyzed using an algorithm for automatic quantification of  
191 microfilariae. The final result was displayed through an app in less than two minutes. The  
192 device showed high sensitivity and specificity (see Table 1) compared to manual counts in  
193 thick blood smears from 33 potentially *Loa*-infected patients, suggesting the potential

194 implications of this device in the field screening of the parasite (26). Such smartphone-  
195 assisted video imaging could be applicable for the diagnosis of other blood parasites and  
196 motile parasitic stages in body fluids or excreta.

197 **Smartphone-assisted microfluidic technology.** Due to their high throughput, easy  
198 handling, parallelism and sensitivity, the use of microfluidic lab-on-a-chip devices (LOCDs)  
199 has greatly increased in medical diagnostics (27). Smartphones offer a tremendous potential  
200 for *in vitro* measurements of biochemical reactions in LOCDs. For instance, Stemple et al.  
201 recently introduced a handheld smartphone-assisted LOCD for the detection of a *P.*  
202 *falciparum* specific protein (HRP-2) (27). Anti-HRP-2-conjugated submicrobeads were  
203 mixed with 10% whole blood sample in a microfluidic LOCD. A smartphone was deployed  
204 for illumination of the sample followed by the detection of the scattered light. Using  
205 scattering/absorption characteristics of the sample, the system was able to measure as low as  
206 1 pg/mL of HRP-2 from blood in 10 min (27).

207 In another study, Liu et al. described an integrated microfluidic chip with a smartphone  
208 recorder for the identification of *Anopheles* spp. (28). The microfluidic device allowed DNA  
209 extraction followed by target DNA amplification using loop mediated isothermal  
210 amplification (LAMP). The amplified products were excited with a DNA intercalating dye  
211 and the fluorescence signal was detected with a smartphone camera. This multiplex system  
212 could be used for parallel identification of several mosquito species. Such a sophisticated  
213 smartphone-based LOCD could be highly useful not only in the onsite diagnosis of parasites  
214 but also in the quick recording of the results and the geographic location of the test for quality  
215 control.

216

## 217 **CONCLUSIONS AND FUTURE IMPLICATIONS**

218 Smartphone microscopy is one of the most common applications of smartphones for the  
219 diagnosis of parasitic diseases. A smartphone allows the direct transfer of images (with  
220 Multimedia Messaging Service (MMS), Bluetooth etc.) to a reference laboratory for quick  
221 assessment, feedback and quality assurance by an expert parasitologist (5, 29). Traditionally,  
222 the Kato-Katz is the commonly used method for the diagnosis of intestinal helminths.  
223 However, it involves laborious manual microscopy and the hookworm ova are rapidly cleared  
224 in this method, resulting in false-negative diagnosis. As an alternative, an inexpensive and  
225 portable ball-lens-mounted smartphone microscope presents a simple POC tool for the  
226 identification of STH in community surveys (10, 11). However, the use of this device is  
227 limited due to various issues such as specimen orientation, hygiene, manual slide navigation,  
228 low sensitivity and small FOV (see Table 2). Some of these issues have been addressed by  
229 applying other lens settings such as a reversed lens (12) and a double-convex lens (13),  
230 though these devices require field-validation.

231 Smartphone-assisted Foldscope and CellScope present attractive POC tools for the  
232 diagnosis of schistosomes as they have been tested in the field, and are lighter in weight and  
233 cost less than US\$1 and US\$6, respectively (14). Despite their high specificity, a major  
234 limitation of these devices is their low sensitivity. This could be explained by their small  
235 FOV (3) or irregular distribution of *Schistosoma* eggs in the excreta. To detect *Schistosoma*  
236 eggs in large field-surveys with improved sensitivity, these devices could be trialed in  
237 conjunction with a specialized algorithm for an automated identification (21). The Newton  
238 NM1 microscope had higher sensitivities than the Foldscope or CellScope for the diagnosis  
239 of *Schistosoma* (15). A combination of the NM1 with the smartphone-algorithm method (21)  
240 could further enhance the sensitivity of this device for field diagnosis of schistosomes.  
241 However, the NM1 could be much more expensive than the Foldscope or CellScope  
242 microscopes. Another way to enhance the sensitivity of smartphone-assisted microscopic

243 devices could be the use of fluorescent-labelled egg binding dyes which produces superior  
244 results than the commonly used McMaster method for the identification of strongyle eggs  
245 (22). A similar system could be trialed for the diagnosis of human helminths. The recent  
246 application of smartphone microscopy for the detection of pathogenic DNA (25, 28) presents  
247 a multiplex potential for parallel identification of several parasitic species in a high-  
248 throughput and short-time format.

249

250 Smartphone-assisted video microscopy is a recent advancement in parasite diagnostics.  
251 For instance, the CellScope Loa allowed the quantification of *L. loa* microfilariae in less than  
252 two minutes with a high sensitivity (100%) (26). Such a device could potentially be applied  
253 for rapid field-diagnosis of other blood-borne parasites such as *Leishmania* and  
254 *Trypanosoma*. Furthermore, video microscopes have the ability to characterize motility  
255 patterns of parasites which could be applied for the diagnosis of flagellate parasites and  
256 parasitic larval stages (30).

257 One of the advantages for using smartphone-based diagnostic tools is the use of dedicated  
258 algorithms and softwares for automated identification of parasites. For instance, the  
259 utilization of a specialized algorithm facilitated smartphone-assisted automatic detection of  
260 *G. lamblia* in large volumes of water in only one hour compared to the conventional methods  
261 which may take one to two days (23). Since water-borne parasitic diseases remain the second  
262 leading cause of death in children under five in developing countries, this technology could  
263 be applicable for large-scale water testing in these regions. Similar to pattern-recognition  
264 algorithms used for human face recognition in the biometric analysis, algorithms could be  
265 developed for the identification of parasites and parasitic eggs. The introduction of pattern-  
266 recognition algorithms for the identification of *Schistosoma* eggs and *P. falciparum* is a  
267 recent advancement in this context (21, 24). Smartphone apps are also setting their feet in

268 parasite diagnostics, for instance TickID (9). Similar apps are required for diagnosis and self-  
269 management of other parasitic infections, especially in resource-constrained regions where  
270 people may own a smartphone despite the inadequacy of basic healthcare facilities (3, 18).  
271 An ideal diagnostic app should work both for iOS and Android systems. Developing  
272 dedicated algorithms and free-downloadable apps for automated diagnosis of parasites offers  
273 a great potential for future parasitology research.

274 Smartphone-assisted microfluidic LOCDs have the potential for high throughput diagnosis  
275 of parasites. For instance, a smartphone-assisted LOCD enabled the detection of *P.*  
276 *falciparum* from whole blood in ~10 min compared to the conventional blood smear method  
277 which may take 1-3 hr (27). Such devices require testing in large-scale field trials for the  
278 diagnosis of important blood-borne parasites. Despite the high robustness of microfluidic  
279 LOCD, they could be more expensive than the conventional diagnostic tools that warrant the  
280 need for studies exploring ways to reduce the cost of such devices for rapid processing a large  
281 number of samples.

282 Most smartphone-based diagnostic devices have been tested in well-controlled laboratory  
283 conditions and for tropical parasitic diseases only. Future studies are required to explore the  
284 usefulness of such devices for the diagnosis of other important parasitic diseases in field  
285 conditions and on clinical specimens. Despite the portability of smartphone-based diagnostic  
286 tools, issues such as manual processing of samples and preparation of microscopic slides  
287 remain to be addressed. Limited battery capacity of smartphones is a major bottleneck for  
288 their field deployability in remote healthcare facilities which can be solved by applying  
289 mobile charging devices with a car-battery or solar-power (29). Internet prices could be high  
290 in low-income regions that may hinder its use in transferring the diagnostic data to a  
291 reference laboratory. The lack of awareness and tangible commercial market are the other  
292 major challenges for smartphone-based diagnostic devices that could be addressed through

293 integrated training, and practical business plans. Sustained research and strong collaboration  
294 among researchers, clinicians, and public sector are required in this context. Currently, there  
295 are no set standards and regulatory approval methods in place for commercialization of  
296 smartphone-based diagnostic devices, which warrants the urgent need for developing  
297 standard guidelines by professional associations/societies such as the World Federation of  
298 Parasitology, the World Association for the Advancement of Veterinary Parasitology and the  
299 American Society for Microbiology. Moreover, these technologies require rigorous quality  
300 control and adequate field validation before deploying them in clinics. A consortium of  
301 experts could be of great help for quality assurance and enhanced usability of such  
302 technologies. Despite all the challenges, these devices have the technical capacity to meet the  
303 enormous diagnostic needs of developing countries with high prevalence of parasitic  
304 diseases.

305 The combined use of smartphones with the inexpensive handheld microscopes and the  
306 microfluidic devices offer a great opportunity for the advancement of portable diagnostic  
307 technologies to overcome the burden of many parasitic diseases in resource-constrained  
308 regions and offer researchers opportunities to develop similar technologies at affordable  
309 prices. Despite lower sensitivity than an established laboratory test, a smartphone-assisted  
310 onsite diagnostic test could be more useful to provide onsite ‘sample-to-answer’ treatment to  
311 the patient infected with a parasitic disease as this patient may not return to the clinic for the  
312 results of a laboratory test (1, 3). The future of smartphone-assisted diagnostic technologies is  
313 very promising and the widespread adoption of such technologies is anticipated in near future  
314 for the accurate and rapid diagnosis of parasitic diseases in an easy-to-use format.

315

316 **SUMMARY**

317 In this mini-review, we have described various smartphone-based devices, applied  
318 specifically for the diagnosis of parasitic diseases of medical and veterinary importance, and  
319 discussed challenges, potential implications and the needs for future development.  
320 Smartphones have been used as a standalone tool or in combination with other microscopic  
321 and microfluidic devices for the identification of various stages of parasites such as eggs,  
322 cysts, and microfilariae. When used with a dedicated algorithm, app or software, smartphones  
323 allow automated and rapid diagnosis of parasites that make them a powerful tool in large  
324 field surveys for disease surveillance and outbreak containment. A major strength of  
325 smartphone-based microscopic devices is their low-cost, widespread availability and onsite  
326 diagnostic potential which could be highly applicable in resource-constrained regions for  
327 effective management of a parasitic disease. As an emerging technology, smartphone-based  
328 diagnostic devices face challenges such as lack of set standards, guidelines, awareness and  
329 tangible commercial market. Despite these challenges, these devices hold the potential to  
330 fulfill the enormous diagnostic needs of developing countries with high prevalence of  
331 parasitic diseases. Future studies are warranted to explore the usefulness of such devices,  
332 especially in field conditions and in clinical settings not only for parasitic but also for other  
333 microbial diseases. The widespread adoption of smartphone-based diagnostic devices is  
334 anticipated in the near future for rapid diagnosis of parasitic diseases.

335

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339

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**Table 1.** Smartphone-based devices for the diagnosis of parasites

| Category                                 | Parasite                                    | Stage                       | Study location | Study type | Sample size | Sensitivity (%)*<br>or Detection<br>limit | Specificity (%)*    | Ref  |      |
|--|---|-----------------------------|----------------|------------|-------------|---|---------------------|------|------|
| Smartphone standalone technology         | <i>Taenia solium</i>                        | Cyst                        | India          | Field      | 1           | -   | -                   | (6)  |      |
|  | <i>Plasmodium falciparum</i>                | -                           | USA            | Lab        | -           | 21 parasites/ $\mu$ L                     | -                   | (7)  |      |
|  | <i>P. falciparum</i>                        | -                           | USA            | Lab        | -           | 1 nM (543 parasites / $\mu$ L)            | -                   | (8)  |      |
|  | Human ticks                                 | -                           | USA            | -          | -           | -   | -                   | (9)  |      |
| Lens-mounted smartphone microscopy       | <i>Ascaris lumbricoides</i>                 | Egg                         | Tanzania       | Field      | 199         | 81  | 87                  | (10) |      |
|  | Hookworm                                    | Egg                         |                |            |             | 14  | 89                  |      |      |
|  | <i>Trichuris trichiura</i>                  | Egg                         |                |            |             | 54  | 63                  |      |      |
|  | <i>T. trichiura</i>                         | Egg                         | Côte d'Ivoire  | Field      | 164, 180    | 31  | 71                  | (11) |      |
|  | <i>Schistosoma mansoni</i>                  | Egg                         |                |            |             | 68  | 64                  |      |      |
| Smartphone-assisted manual microscopy    | <i>A. lumbricoides</i>                      | Egg                         | USA            | Lab        | -           | -   | -                   | (12) |      |
|  | <i>A. lumbricoides</i>                      | Egg                         | New Zealand    | Lab        | -           | -   | -                   | (13) |      |
|  | <i>S. haematobium</i>                       | Egg                         | Ghana          | Field      | 49          | 56  | 93                  | (14) |      |
|  | <i>S. haematobium</i>                       | Egg                         |                |            |             | 68  | 100                 |      |      |
|  | <i>S. haematobium</i>                       | Egg                         | Côte d'Ivoire  | Field      | 226         | 36  | 100                 | (15) |      |
|  | <i>S. mansoni</i>                           | Egg                         |                |            |             | 50  | 100                 |      |      |
|  | <i>Giardia lamblia</i>                      | Cyst                        | USA            | Lab        | -           | -   | -                   | (16) |      |
|  | <i>G. lamblia</i>                           | Cyst                        | USA            | Lab        | -           | -   | -                   | (17) |      |
|  | <i>P. falciparum</i>                        | -                           | USA            | Lab        | -           | -   | -                   | (18) |      |
|  | <i>P. falciparum</i>                        | -                           | Côte d'Ivoire  | Field      | 223         | 80  | 100                 | (19) |      |
| Smartphone-assisted automated microscopy | <i>P. falciparum</i>                        | Mixed                       | Uganda         | Lab        | -           | -   | -                   | (5)  |      |
|  | <i>P. chabaudi</i>                          | Hemozoin                    | USA            | Lab        | -           | -   | -                   | (20) |      |
|  | <i>S. haematobium</i>                       | Egg                         | Sweden         | Lab        | -           | 79  | 100                 | (21) |      |
|  | Animal parasites                            | Egg                         | USA            | Lab        | -           | -   | -                   | (22) |      |
|  | <i>G. lamblia</i>                           | Cyst                        | USA            | Lab        | -           | 12 cysts /10 mL                           | 94 (50 cysts/10 mL) | (23) |      |
|  | <i>P. falciparum</i>                        | Trophozoite                 | Portugal       | Lab        | 6           | 81  | 94                  | (24) |      |
|  | <i>Trypanosoma cruzi</i>                    | DNA                         | USA            | Lab        | -           | -   | -                   | (25) |      |
|  | <i>Loa loa</i>                              | Microfilaria                | Cameroon       | Field      | 33          | 100                                       | 94                  | (26) |      |
|  | Smartphone assisted microfluidic technology | <i>P. falciparum</i>        | -              | USA        | Lab         | -   | 1 pg/mL             | -    | (27) |
|  |   | <i>Anopheles arabiensis</i> | DNA            | USA        | Lab         | -   | -                   | -    | (28) |
| <i>An. gambiae</i>                       |   | DNA                         |                |            |             | -   | -                   |      |      |

- not available/applicable; \* Compared to a conventional method, e.g. a standard microscope or manual identification; decimal points of these figures were rounded off.

**Table 2.** A comparison of smartphone-based devices in the diagnosis of parasites

| Category  | Main device/<br>technology                                | Advantages   | Limitations  | Ref      |
|---|---|--|--|----------|
| <b>Smartphone standalone technology</b>         | iPhone  | <ul style="list-style-type: none"> <li>◦ Rapid</li> <li>◦ Avoids false-negative diagnosis</li> </ul>   | <ul style="list-style-type: none"> <li>◦ Validated with one study only</li> </ul>  | (6)      |
|   | iPhone 5S + algorithm                                     | <ul style="list-style-type: none"> <li>◦ Automated</li> <li>◦ Reduced reporting errors</li> <li>◦ Avoids false-negative diagnosis</li> </ul>         | <ul style="list-style-type: none"> <li>◦ Lower sensitivity than visual method</li> </ul>   | (7)      |
|   | iPhone 6S + smartphone application (app)                  | <ul style="list-style-type: none"> <li>◦ Automated</li> <li>◦ Reduced reporting errors</li> <li>◦ Avoids false-negative diagnosis</li> </ul>         | <ul style="list-style-type: none"> <li>◦ Lower sensitivity than visual method</li> </ul>   | (8)      |
| <b>Lens-mounted smartphone microscopy</b>       | iPhone 4s + ball-lens                                     | <ul style="list-style-type: none"> <li>◦ Low-cost</li> <li>◦ Portable</li> </ul>   | <ul style="list-style-type: none"> <li>◦ Small field-of-view (FOV)</li> <li>◦ Low sensitivity</li> </ul>                                   | (10, 11) |
|   | iPhone 4S + reversed camera lens                          | <ul style="list-style-type: none"> <li>◦ Low-cost</li> <li>◦ Portable</li> <li>◦ Relatively larger FOV</li> </ul>                                    | <ul style="list-style-type: none"> <li>◦ Illumination and vignetting issues</li> <li>◦ Field validation required</li> </ul>                | (12)     |
|   | Nokia Lumia 1020 + double convex objective lens + ImageJ  | <ul style="list-style-type: none"> <li>◦ Composite imaging of eggs scattered on different focal planes</li> </ul>                                    | <ul style="list-style-type: none"> <li>◦ Lower resolution</li> <li>◦ Field validation required</li> </ul>                                  | (13)     |
| <b>Smartphone-assisted manual microscopy</b>    | iPhone 5S + Foldscope                                     | <ul style="list-style-type: none"> <li>◦ Low-cost</li> <li>◦ Portable</li> <li>◦ High specificity</li> </ul>   | <ul style="list-style-type: none"> <li>◦ Limited sensitivity</li> <li>◦ Manual slide navigation issue</li> </ul>                           | (14)     |
|   | iPhone 5S + CellScope                                     | <ul style="list-style-type: none"> <li>◦ Low-cost</li> <li>◦ Portable</li> <li>◦ High specificity</li> </ul>   | <ul style="list-style-type: none"> <li>◦ Low sensitivity</li> <li>◦ Manual slide-navigation issue</li> </ul>                               | (14, 15) |
|   | Motorola ZN5 + LED + aperture                             | <ul style="list-style-type: none"> <li>◦ Lensfree and Light-weight</li> <li>◦ Avoids undesired artifacts caused by demosaicing algorithms</li> </ul> | <ul style="list-style-type: none"> <li>◦ Powerful smartphone required to image holograms</li> <li>◦ Field validation required</li> </ul>   | (16)     |
|   | Sony-Erickson U10i Aino + external lens + color filter    | <ul style="list-style-type: none"> <li>◦ Fluorescent microscopy</li> <li>◦ Large FOV</li> <li>◦ Long depth-of-field</li> </ul>                       | <ul style="list-style-type: none"> <li>◦ Field validation required</li> </ul>  | (17)     |
|   | Nokia N73 + lens assembly                                 | <ul style="list-style-type: none"> <li>◦ Wide-field imaging</li> <li>◦ Allows both bright-field and fluorescent imaging</li> </ul>                   | <ul style="list-style-type: none"> <li>◦ Field validation required</li> </ul>  | (18)     |
|   | iPhone 5S + Newton NMI microscope                         | <ul style="list-style-type: none"> <li>◦ Handheld</li> <li>◦ Portable</li> <li>◦ High specificity</li> </ul>   | <ul style="list-style-type: none"> <li>◦ Limited sensitivity</li> <li>◦ Relatively expensive</li> <li>◦ Slide-navigation issues</li> </ul> | (19)     |
|   | Nokia candy bar models + light microscope                 | <ul style="list-style-type: none"> <li>◦ High quality imaging</li> <li>◦ Image sharing facility</li> </ul>   | <ul style="list-style-type: none"> <li>◦ Heavy microscope</li> </ul>   | (5)      |
|   | iPhone 5S + polarized microscopic assembly                | <ul style="list-style-type: none"> <li>◦ Improved contrast</li> <li>◦ Time-saving</li> <li>◦ Operable by less-skilled personnel</li> </ul>           | <ul style="list-style-type: none"> <li>◦ Field validation required</li> <li>◦ High-resolution lens required</li> </ul>                     | (20)     |
| <b>Smartphone-assisted automated microscopy</b> | Nokia E71 or Sony Ericsson C905 + algorithms              | <ul style="list-style-type: none"> <li>◦ Automated</li> <li>◦ Time-saving</li> <li>◦ High specificity</li> </ul>                                     | <ul style="list-style-type: none"> <li>◦ Images from each device needs separate validation</li> </ul>                                      | (21)     |
|   | iPhone 5s or Sony Experia Z3 or Nokia Lumia 1020 + ImageJ | <ul style="list-style-type: none"> <li>◦ Automated</li> <li>◦ Superior results than McMaster method</li> </ul>                                       | <ul style="list-style-type: none"> <li>◦ Field validation required</li> <li>◦ Fluorescent-labelling required</li> </ul>                    | (22)     |
|   | Nokia Lumia 1020 + algorithm                              | <ul style="list-style-type: none"> <li>◦ Large FOV and sample load</li> <li>◦ Automated</li> <li>◦ Portable</li> </ul>                               | <ul style="list-style-type: none"> <li>◦ Field-validation required</li> </ul>  | (23)     |
|   | HTC 1S or LG Nexus 5 + algorithm + app                    | <ul style="list-style-type: none"> <li>◦ Automated</li> <li>◦ High specificity</li> </ul>  | <ul style="list-style-type: none"> <li>◦ Limited sensitivity</li> <li>◦ Pre-annotation of images could be challenging</li> </ul>           | (24)     |
|   | Samsung Galaxy S + thermal cycler + MATLAB + ImageJ       | <ul style="list-style-type: none"> <li>◦ Automated</li> <li>◦ No need for sample preparation</li> </ul>  | <ul style="list-style-type: none"> <li>◦ Complex data analysis</li> <li>◦ Thermal cycler required</li> </ul>                               | (25)     |
|   | iPhone 5S + reversed                                      | <ul style="list-style-type: none"> <li>◦ Automated video</li> </ul>  | <ul style="list-style-type: none"> <li>◦ Relatively complex</li> </ul>   | (26)     |

|  |   |  |  |      |
|--|---|--|--|------|
|  | lens cell scope +<br>algorithm + app                | <ul style="list-style-type: none"> <li>◦ High sensitivity and specificity</li> <li>◦ No need for sample preparation</li> </ul> | imaging<br>design  |      |
| <b>Smartphone assisted<br/>microfluidic technology</b> | iPhone 4 + Optofluidic<br>device + Photoshop<br>CS5 | <ul style="list-style-type: none"> <li>◦ Rapid</li> <li>◦ Portable</li> <li>◦ High sensitivity</li> </ul>                      | <ul style="list-style-type: none"> <li>◦ Expertise required for data analysis</li> <li>◦ Field validation required</li> <li>◦ Fluorescence-labelling required</li> </ul> | (27) |
|  | iPhone 4 + microfluidic<br>device                   | <ul style="list-style-type: none"> <li>◦ Rapid</li> <li>◦ Portable</li> <li>◦ Small sample size required</li> </ul>            | <ul style="list-style-type: none"> <li>◦ Field validation required</li> <li>◦ DNA amplification required</li> <li>◦ Relatively expensive</li> </ul>                      | (28) |