Differential Diagnosis Assessment in Ambulatory Care With an Automated Medical History–Taking Device: Pilot Randomized Controlled Trial

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Abstract

Background: Automated medical history–taking devices (AMHTDs) are emerging tools with the potential to increase the quality of medical consultations by providing physicians with an exhaustive, high-quality, standardized anamnesis and differential diagnosis.

Objective: This study aimed to assess the effectiveness of an AMHTD to obtain an accurate differential diagnosis in an outpatient service.

Methods: We conducted a pilot randomized controlled trial involving 59 patients presenting to an emergency outpatient unit and suffering from various conditions affecting the limbs, the back, and the chest wall. Resident physicians were randomized into 2 groups, one assisted by the AMHTD and one without access to the device. For each patient, physicians were asked to establish an exhaustive differential diagnosis based on the anamnesis and clinical examination. In the intervention group, residents read the AMHTD report before performing the anamnesis. In both the groups, a senior physician had to establish a differential diagnosis, considered as the gold standard, independent of the resident’s opinion and AMHTD report.

Results: A total of 29 patients were included in the intervention group and 30 in the control group. Differential diagnosis accuracy was higher in the intervention group (mean 75%, SD 26%) than in the control group (mean 59%, SD 31%; P=.01). Subgroup analysis showed a between-group difference of 3% (83% [17/21]-80% [14/17]) for low complexity cases (1-2 differential diagnoses possible) in favor of the AMHTD (P=.76), 31% (87% [13/15]-56% [18/33]) for intermediate complexity (3 differential diagnoses; P=.02), and 24% (63% [34/54]-39% [14/35]) for high complexity (4-5 differential diagnoses; P=.08). Physicians in the intervention group (mean 4.3, SD 2) had more years of clinical practice compared with the control group (mean 5.5, SD 2; P=.03). Differential diagnosis accuracy was negatively correlated to case complexity (r=0.41; P=.001) and the residents’ years of practice (r=0.04; P=.72). The AMHTD was able to determine 73% (SD 30%) of correct differential diagnoses. Patient satisfaction was good (4.3/5), and 26 of 29 patients (90%) considered that they were able to accurately describe their symptomatology. In 8 of 29 cases (28%), residents considered that the AMHTD helped to establish the differential diagnosis.

Conclusions: The AMHTD allowed physicians to make more accurate differential diagnoses, particularly in complex cases. This could be explained not only by the ability of the AMHTD to make the right diagnoses, but also by the exhaustive anamnesis provided.
We tested a novel AMHTD, named DIAANA (DIAgnosis & ANAmneses; created by Logic-based Medicine Sàrl), to help the physician to establish the differential diagnosis (DD) more accurately, based on broad possibilities of disease or trauma localization, triggering factors, and symptoms. The physician can therefore begin his consultation with an exhaustive anamnesis summary including a more precise localization and nature of symptoms as well as a high-sensitivity DD list with corresponding triggering factors for each diagnosis. We consider that this tool could help the physician in his/her diagnostic reasoning and to perform tasks more efficiently, without being substituted by the AMHTD.

**Methods**

**Study Design**

We conducted a pilot, single-center, unblinded, 1:1 parallel-group, randomized efficacy trial. No follow-up was necessary. There were no changes in the protocol after trial commencement. The study protocol was optimized and approved by an independent expert methodologist. It was not registered as it was considered to be a pilot phase. Given that recruitment began just after the approval, it would therefore have not been relevant to register the study after the beginning of the recruitment. The protocol was approved by the Medical Ethics Committee of Geneva University Hospitals (Geneva, Switzerland; REQ-2017-00878). No bugs were fixed during the trial. As this was a purely observational study without identifiable side effects or negative consequences for patients, only oral informed consent was obtained, supported by a brief written description of the project. Consolidated Standards of Reporting Trials of Electronic and Mobile Health Applications and online TeleHealth V 1.6 (see Multimedia Appendix 1) was used to improve and standardize the quality of this paper [25].

**Patient Population**

From May to September 2018, we prospectively enrolled adult patients presenting to the emergency outpatient unit of our institution and suffering from symptoms covered by the AMHTD. Symptoms were localized to the superior member (apart from the hand, as the device had not yet been programmed to take related conditions into consideration), the trunk, and the inferior member, with the exception of strictly dermatologic concerns and toes and inversion ankle trauma as the diagnosis is generally obvious. We excluded patients with a medical situation considered as urgent and unable to complete the digitalized AMHTD (sight problems, advanced age, and non-French-speaking). Patients were enrolled only when one of the senior physicians in charge of the project (CL, TW, RG, MB, and HS) and one of the coordinators (CJ and BV) were present.

**Objectives**

The primary aim of this pilot study was to investigate whether the DIAANA AMHTD allowed physicians to establish a more accurate DD, with the DD of a senior physician considered as the gold standard. Secondary aims were to assess the accuracy of the DD list established by the AMHTD, identify factors that might influence the usefulness of the AMHTD, and evaluate physician and patient satisfaction with its use.
Randomization and Recruitment
At the beginning of the study, 18 residents of the emergency outpatient unit were stratified, and 1:1 matched by their years of clinical experience (orthopedics, rheumatology, and physical medicine counted twice) and then randomized. When a patient was allocated to a resident physician using the emergency software system, the coordinating researcher evaluated the patient’s potential eligibility. The senior physician then confirmed the patient’s eligibility and applied the exclusion criteria. Depending on the resident physician’s allocation, the patient was included in either the intervention or the control group. In each group, the recruitment was blocked after the inclusion of 30 patients.

DIAANA Tool Presentation
The DIAANA AMHTD functions as follows: On the basis of an interactive questionnaire completed by the patient before the consultation, which includes 269 questions (mainly multiple choice), it performs an exhaustive anamnesis focused on the problem and proposes a panel of DDs with a high sensitivity, selected on a panel of 126 diagnostic entities. The artificial reasoning system of DIAANA mimics how a specialist physician would reason to establish a DD. The information transmitted is in an easy-to-use form for the physician that includes a summary of the anamnesis centered on relevant elements from the questionnaire and a list of possible diagnoses with their emergency level, potential contributing factors, and first-line management proposals. Multimedia Appendix 2 illustrates an example of a patient suffering from deep vein thrombosis that was initially confounded with a tennis leg. More detailed information is available on the AMHTD’s website [26].

DIAANA Tool Development
For 3 years, AS was involved in the development of the AMHTD, taking into consideration all aspects of the diagnosis and management of orthopedic, rheumatologic, vascular, neuropathic, and sports-related medical conditions, with the help of a few sources [27-29] as well as peer advice.

The system was built with triggering conditions that are turned on when the patient selects a specific answer. The triggering condition will then call up new questions and diagnostic entities. As an example, if the patient clicks leg on the general localization, the trigger leg is turned on, and a more specific question about the leg localization appears (see Multimedia Appendix 2). AS built a first draft of DIANNA including the principal questions of a proper musculoskeletal anamnesis. Then, he considered the 126 selected diagnosis entities in more depth and added more specific questions for each diagnosis step by step. The accuracy of DIANNA depends, therefore, on the accuracy of the patient’s answer as well as the exhaustivity of the questions and diagnostic entities. As an example, if the correct localization (eg, ankle) is not selected, specific questions (eg, trauma in external rotation) and a specific diagnosis (eg, syndesmosis sprain) will not be triggered and thus be missing in the DIANNA summary.

Hundreds of episodes of testing with healthy volunteers, medical students, and patients were performed during the development process, and the formulation of questions, triggering conditions, and the DIANNA summary were adjusted according to feedback from users. A final development phase was conducted with the feedback of 20 patients presenting to the emergency outpatient unit, and the first version of the digital content of the tool was then frozen for the pilot study. This frozen version remains available upon request to the corresponding author.

Intervention
In the intervention group, patients in the AMHTD group were asked to complete a digital form on a touch pad by the coordinator (and without help) before the medical consultation. The AMHTD summary was then printed and given to the resident physician before the consultation. At the end of the consultation, but before consulting the complementary medical examination results (radiographs and blood laboratory results), the resident physician established his/her DD on the diagnosis list (see Multimedia Appendix 3) on a touch pad, without the help of the research coordinator. In parallel, the senior physician established the gold standard DD on the same list. In the control group, the resident physician established his DD on the diagnosis list at the end of the consultation, but before consulting the complementary medical examinations. The senior physician followed the same procedure. For ethical reasons, the use of the AMHTD had no influence on patient care as the clinical management was fully decided upon by the senior physician who had no access to the summary generated.

Outcomes
The primary outcome was the percentage of correct DDs established by the resident physician compared with the senior physician. Secondary outcomes included (1) the percentage of correct AMHTD DDs and the percentage of correct AMHTD DDs followed/not followed by the resident, as well as the percentage of incorrect AMHTD DDs followed by the resident and the number of incorrect AMHTD DDs; (2) overall patient satisfaction on the understandability of AMHTD questions (1-5 Likert scale), ability to describe symptoms accurately (percentage), and respect of the patient’s wish to use the AMHTD at home and to keep the generated summary; (3) resident’s feedback on the wish to obtain the AMHTD summary; (4) overall patient satisfaction; (5) resident’s feedback on the wish to keep the generated summary (percentage); and (6) the number of incorrect AMHTD DDs that would have been omitted otherwise (percentage), and if the use of the device saved time (1-5 Likert scale); and (4) the percentage of correct DDs depending on case complexity, defined as the number of DDs present in the gold standard DD (1-2 DDs=low complexity; 3 DDs=intermediate complexity; and 4-5 DDs=high complexity). The stratification for the case complexity definition used has never been published. The rationale was to highlight that the AMHTD was built and conceived to help the physician when the diagnosis might be confusing or in the case of a complex situation. Indeed, it would not be relevant to ask the patient to provide a complete anamnesis if the physician can complete it in 2 min for a problem such as benign soft tissue trauma.

Statistical Analyses
A sample size of 30 patients per group was chosen as recommended for pilot studies to achieve an appropriate level of statistical power [30]. It corresponds to the detection of a...
potential difference of 21% between groups for a power of 80% and an alpha significance level of 5%. Descriptive statistics were used to describe baseline characteristics. Differences between groups in the intention-to-treat analysis were evaluated using Student t test or the Wilcoxon rank-sum test, when appropriate. Analysis of covariance was performed considering the covariables of interest (primary outcome, case complexity, and resident’s years of experience) with a P value < .20 considered as significant in univariate analysis. P values < .05 were considered as statistically significant. All analyses were performed using R v3.4.2 Portable (Free Software Foundation Inc).

Results

Population

Of the 81 patients screened, 64 were randomized and allocated to residents (Figure 1). Among the randomized patients, 4 allocated to the intervention group were not included as 30 patients were already included in the intervention group; 1 patient was lost to follow-up. In the final analysis, 29 patients were included in the intervention group and 30 in the control group. Preintervention patient demographics, case complexity, and initial complaint/s did not differ between the groups (Table 1). Residents in the control group had more years of practice (P=.03).

Figure 1. Study flow chart. AMHTD: automated medical history-taking device; DD: differential diagnosis.
Table 1. Baseline characteristics.

| Baseline characteristics                  | AMHTD<sup>a</sup> (n=29) | Control group (n=30) | \( P \) value |
|------------------------------------------|---------------------------|----------------------|---------------|
| **Age (years)**                          |                           |                      |               |
| Mean (SD)                                | 38 (14)                   | 42.1 (16)            | .29           |
| Range                                    | 17-66                     | 19-75                | .29           |
| **Male gender, n (%)**                   |                           |                      |               |
|                                         | 23 (79)                   | 22 (73)              | .82           |
| **Physician’s practice (years)**         |                           |                      |               |
| Mean (SD)                                | 4.3 (2)                   | 5.5 (2)              | .03           |
| Range                                    | 3-8                       | 3-8                  | .03           |
| **Case complexity (number of differential diagnoses to find)** |               |                      |               |
| Mean (SD)                                | 3.1 (1)                   | 2.9 (1)              | .60           |
| Range                                    | 1-5                       | 1-5                  | .60           |
| **Initial complaint, n (%)**             |                           |                      |               |
| Elbow pain                               | 1 (3)                     | 1 (3)                | >.99          |
| Shoulder pain and trauma                 | 3 (10)                    | 2 (7)                | .97           |
| Back pain and trauma                     | 5 (17)                    | 7 (23)               | .80           |
| Pelvic pain                              | 2 (7)                     | 0 (0)                | .46           |
| Knee pain and trauma                     | 8 (28)                    | 6 (20)               | .70           |
| Ankle trauma                             | 4 (14)                    | 6 (20)               | .77           |
| Foot trauma                              | 2 (7)                     | 2 (7)                | >.99          |
| Soft tissue trauma and swelling          | 4 (14)                    | 5 (17)               | >.99          |

<sup>a</sup>AMHTD: automated medical history–taking device.

Analysis of Accuracy of Differential Diagnosis

In the univariate analysis, the percentage of correct DDs was (1) higher in the intervention group (mean 75% [SD 26%] vs mean 59% [SD 31%], respectively; \( P = .03 \)); (2) negatively correlated to case complexity (\( r = 0.41; P = .001 \)); and (3) negatively correlated to residents’ years of practice (\( r = 0.04; P = .72 \)). The \( P \) value of the analysis of covariance model, including the percentage of DDs found and case complexity was .01. Considering case complexity, we observed between-group differences in favor of the AMHTD of 3% (83% [17/21]-80% [14/17]) for low-complexity cases, 31% (87% [13/15]-56% [18/33]) for intermediate-complexity cases, and 24% (63% [34/54]-39% [14/35]) for high-complexity cases (Table 2). The type of DD made by the senior physician, depending on the case complexity, is presented in the Multimedia Appendix 4.

By comparison, the AMHTD was able to find 73% (SD 30%) of correct DDs for the whole cohort: 91% (SD 20%) for low-complexity cases; 67% (SD 24%) for moderate-complexity cases; and 58% (SD 32%) for high-complexity cases (see Multimedia Appendix 5). The AMHTD also proposed 5(SD 4) incorrect diagnostic proposals. Residents did not list 10% (SD 19%) of the correct DDs proposed by the AMHTD and listed 21% (SD 51%) of incorrect DDs.

Table 2. Percentage of correct differential diagnoses per group.

| DD<sup>a</sup> studied | AMHTD<sup>b</sup> (n=29) | Control group (n=30) | Univariate analysis \( P \) value | Multivariate analysis \( P \) value |
|-------------------------|---------------------------|----------------------|----------------------------------|-----------------------------------|
| Mean (SD)               | Range                     | Mean (SD)            | Range                           |                                   |
| DD accuracy             | 75 (26)                   | 25-100               | 59 (31)                          | 0-100                             | .03                               | <.001                             |
| Low complexity (1-2 DDs to find) | 83 (25) | 50-100               | 80 (26)                          | 50-100                            | .76                               | __<sup>c</sup>                      |
| Moderate complexity (3 DDs to find) | 87 (18) | 67-100               | 56 (26)                          | 0-100                             | .02                               | __                                |
| High complexity (4-5 DDs to find) | 63 (25) | 25-100               | 39 (29)                          | 0-80                              | .08                               | __                                |

<sup>a</sup>DD: differential diagnosis.

<sup>b</sup>AMHTD: automated medical history–taking device.

<sup>c</sup>Not applicable.
Users Satisfaction
Patient satisfaction was good regarding overall satisfaction with questions and their understandability, and 26 of 29 (90%) patients considered that they were able to accurately describe their symptoms. Of note, 14 of 29 (48%) patients wished to use the AMHTD at home, and 20 of 29 (69%) resident physicians wished to obtain the full report of the AMHTD. Although 8 of 29 (28%) residents considered that the device helped to establish the DD, they estimated overall that the AMHTD was neither time-saving nor time-wasting (see Multimedia Appendix 6).

Discussion
Principal Findings
Our results confirmed that the AMHTD significantly allowed the physician to establish a more exhaustive DD (from 59% to 75%). This effect was more important in moderate-complexity (from 56% to 87%) and high-complexity (from 39% to 63%) cases. Of note, the diagnostic list established by the AMHTD was not as accurate as expected (73%, 66/90) and was more precise for low-complexity cases. Overall patient satisfaction (4.3/5) was good, including the ability to accurately describe the presented symptomatology (90%, 26/29). Thus, our results were in agreement with the main factors that guarantee the success of electronic health (eHealth) [31], that is, an improved diagnosis and clinical management, as well as patient-centered care. Our panel of patients presenting to the outpatient unit had common pathologies and was managed by residents at the end of their training. These conditions are common in outpatient services in Switzerland, and our results should be applicable to other hospitals in the country.

Limitations
Our study has some limitations. First, it was an unblinded pilot study with a limited sample size in 1 care center. Therefore, we did not anticipate statistically significant results and did not register our protocol following ethics committee approval. Second, our groups were not balanced as resident physicians in the control group had more years of practice, thus leading to a potential selection bias that could have induced an overestimation of the ability to find a correct DD in the control group. Therefore, the positive effect of 16% (75% [68/90]-59% [50/85]) on the accuracy of the DD might be underestimated. Third, even though our senior physicians were experts in the fields of orthopedics and emergency medicine, the gold standard DD might be flawed, especially in more complex cases. This may be a potential explanation for the observed poorer accuracy of the AMHTD DDs in complex cases. Finally, our AMHTD is still under development, and the reliability of patient responses may be suboptimal, especially because of the absence of images to help in patient symptom localization. This could potentially lead to a degree of uncertainty related to the summary generated. Concerning the DIANNA tool digital content, even if we are fully satisfied with the anamnesis summary, the list of diagnoses might lack accuracy.

Interpretation and Comparison With Prior Research
At present, artificial intelligence systems are still unable to replace physicians for the establishment of a correct DD [31]. Despite this, artificial intelligence allows to complement the work of the physician [32] and even establish an accurate list of problems [33] as shown recently with IBM Watson. The physician’s ability to establish a DD can be improved by providing a case summary and a list of possible diagnoses [32,34]. In contrast with other existing digital systems designed to work hand-to-hand with the physician, such as Ada (Ada Health GmbH), K (K Health), and the Mayo Clinic Symptom Checker (Mayo Clinic), DIAANA is focused on the anamnesis rather than the diagnosis, and highly specialized in injury/disease of the musculoskeletal system. To the best of our knowledge, these abovementioned systems have not been challenged in randomized trials. In addition, we were unable to find any relevant literature concerning other similar systems in the field of general medicine or orthopedics. For instance, in the field of psychiatry, a self-report tool allowed the physician to perform a more accurate diagnosis [35]. Similarly, in acute pediatric assessment, it was shown that junior physicians were able to significantly improve the quality of their diagnostic workup and reduce diagnostic omission errors with the use of a Web-based diagnostic reminder system [36]. These observations are concordant with our results as we showed that it was possible to significantly improve the quality of the DD by providing the physician with an exhaustive anamnesis summary and a list of possible DDs. However, in our study, whether the physician was helped by the exhaustive anamnesis summary or by the DD panel remains open. Both may be useful, although we would suggest that the medical history summary may be superior as the DD panel was not as accurate as expected. Indeed, the DD accuracy of the AMHTD alone (73%, 66/90) was slightly superior to the resident physician in the control group (59%, 55/85), but not superior to the resident physician aided by the AMHTD (75%, 68/90). The reliability of the AMHTD DD without the interpretation of the physician is, therefore, not sufficient. On the other hand, the physician may have underestimated the AMHTD DD reliability, as 10% (9/90) of diagnoses were omitted by residents, but suggested by the AMHTD. This means that if the physician had systematically followed the suggestions of the AMHTD, he/she would have found 85% (78/90) of correct DDs instead of 75% (68/90). The physician should be also aware that the correct diagnosis may be absent on the diagnosis list and, in this case, he/she should not waste energy and resources by trying to explore the entire diagnosis list in depth.

The AMHTD presented was conceptualized as a consultation complement for the physician, and not as a substitute. Physician-informatics partnership is the cornerstone of quality of care improvement, not only because it preserves human relationships [31,37], but also because it is the only condition under which diagnostic assistance has been proven to date. In addition to the existing solutions presented above, it has been shown that patients with unresolved medical issues who submitted their cases on the Web to a panel of specialized case-solvers estimated being helped in their diagnosis process in 60% of the cases [38]. We used the DD as a primary outcome rather than the finally retained diagnosis. Even if only the final diagnosis makes clinical sense, it is well known that only an exhaustive DD can lead to a correct diagnosis with any certainty in medical practice. Using the DD as a primary outcome allowed...
to increase the effect size because the success rate in establishing a DD is poorer than finding the correct diagnosis. Moreover, to identify situations where a rare but serious diagnosis is missed, thousands of patients should be included if the primary outcome was to be considered as the final diagnosis.

The use of eHealth devices for training purposes is on the rise, as reflected in the increasing use of anamnesis and diagnostic supporting tools used by medical students [39]. We consider that our AMHTD presents ideal characteristics for the training of resident physicians by providing an exhaustive anamnesis and a list of DDs with their degree of emergency and associated factors, as well as initial management guidance. Moreover, the device could be used as a tool for asynchronous teleconsultation.

Workload and workflow disruption are recognized as negative factors influencing the outcome of eHealth interventions [31]. We hypothesized that the exhaustive information collected by the AMHTD would allow the physicians to gain some time. Surprisingly, our physicians estimated that the AMHTD was neither time-saving nor time-wasting. Unfortunately, it was not possible to differentiate the potential time gain for clinical evaluation and reasoning from the time associated with the study itself, for example, contact with the coordinating researcher or waiting for the AMHTD summary to be generated. It is also possible that in low-complexity cases, where the medical history is easily performed, the AMHTD becomes time-consuming. We were unable to measure objectively the consultation time, which may be fragmented when physicians are managing more than one patient at the same time. Completion of the AMHTD form takes some time for patients (20 min in our experience). However, as evidenced by the high satisfaction rate, patients are generally happy to take the necessary time to complete the form. In our study, patients completed the AMHTD form when the waiting time was estimated to be greater than 20 min before the start of the consultation.

Overall, patient satisfaction was good. Of 29 patients, 12 (41%) expressed willingness to keep the AMHTD at home, thus emphasizing the subjective importance for the patients to keep their medical folder and the eHealth tool. We did not provide patients with the AMHTD summary because of the necessity to remain noninterventional in the context of the study for ethical purposes and to avoid causing anxiety to patients when reading highly sensitive DDs. A minority of residents (8/29, 28%) considered the AMHTD as meaningful, and this might reflect the lack of usefulness of the AMHTD for low-complexity cases. Interestingly, 69% (20/29) of physicians wished to obtain the entire AMHTD form, thus potentially highlighting the need to obtain the most accurate and least transformed information as possible, even to the detriment of their time. This contrasts with our initial point of view that the AMHTD summary was sufficient, and the full form would lead to time loss for the physician.

Conclusions
The tested musculoskeletal-focused AMHTD allowed physicians to make a more accurate DD, particularly for complex cases. This could be explained not only by the ability of the AMHTD to propose the right diagnosis but also by the exhaustive anamnesis provided. Patients and physicians expressed overall satisfaction with the process. On the basis of these pilot study results, further research will aim to assess and clarify the following points: confirmation of the findings and a fine-tuned assessment of the accuracy of the established DD, depending on complexity; objective measurement of consultation time; and an evaluation of the physicians’ learning curve, both in terms of the accuracy of the DD and duration of the consultation.

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Authors’ Contributions
AS, CB, and HS designed the study and provided input throughout the study. CJ, RG, and CL collected the data. HS provided clinical expertise throughout the study and assisted with the finalization of the instrument. CB analyzed the data, assisted by AS and CJ. AS wrote the manuscript together with contributions from all authors. All authors read and approved the final manuscript.

Conflicts of Interest
AS and, to a lesser extent, HS are partners in the limited liability company that owns the DIAANA AMHTD. To decrease any conflicts of interest as much as possible, the choice of the study design and the statistical analyses were the responsibility of CL and CB, with the support of Angèle Gayet-Ageron.

Editorial Notice
This randomized study was not registered. The editor granted an exception of ICMJE rules for prospective registration of randomized trials because the risk of bias appears low and the study was considered formative. However, readers are advised to carefully assess the validity of any potential explicit or implicit claims related to primary outcomes or effectiveness.

Multimedia Appendix 1
CONSORT - EHEALTH checklist (V 1.6.1).
Multimedia Appendix 2

Example of a patient complaining of muscle cramp.

Multimedia Appendix 3

Differential diagnosis list.

Multimedia Appendix 4

Types of differential diagnoses selected by the senior physician for each level of complexity.

Multimedia Appendix 5

Differential diagnoses found by the automated medical history-taking device.

Multimedia Appendix 6

Automated medical history-taking device group: patient and resident physician satisfaction.

References

1. Phillips RL, Bartholomew LA, Dovey SM, Fryer GE, Miyoshi TJ, Green LA. Learning from malpractice claims about negligent, adverse events in primary care in the United States. Qual Saf Health Care 2004 Apr;13(2):121-126 [FREE Full text] [doi: 10.1136/qshc.2003.008029] [Medline: 15069219]

2. Holohan TV, Colestro J, Grippi J, Converse J, Hughes M. Analysis of diagnostic error in paid malpractice claims with substandard care in a large healthcare system. South Med J 2005 Nov;98(11):1083-1087. [doi: 10.1097/01.smj.0000170729.51651.f7] [Medline: 16351028]

3. Sandars J, Esmail A. The frequency and nature of medical error in primary care: understanding the diversity across studies. Fam Pract 2003 Jun;20(3):231-236. [doi: 10.1093/fampra/cmg301] [Medline: 12738689]

4. Chandra A, Nundy S, Seabury SA. The growth of physician medical malpractice payments: evidence from the National Practitioner Data Bank. Health Aff (Millwood) 2005(Suppl Web Exclusives):W5-240. [doi: 10.1377/hlthaff.w5.240] [Medline: 15928255]

5. Thomas EJ, Studdert DM, Newhouse JP, Zbar BI, Howard KM, Williams EJ, et al. Costs of medical injuries in Utah and Colorado. Inquiry 1999;36(3):255-264. [Medline: 10570659]

6. Graber M. Diagnostic errors in medicine: a case of neglect. Jt Comm J Qual Patient Saf 2005 Feb;31(2):106-113. [doi: 10.1016/S1553-7250(05)31015-4] [Medline: 15791770]

7. Singh H, Meyer AN, Thomas EJ. The frequency of diagnostic errors in outpatient care: estimations from three large observational studies involving US adult populations. BMJ Qual Saf 2014 Sep;23(9):727-731 [FREE Full text] [doi: 10.1136/bmjqs-2013-002627] [Medline: 24742777]

8. Schiff G, Kim S, Abrams R, Cosby K, Lambert B, Elstein A, et al. Diagnosing diagnosis errors: lessons from a multi-institutional collaborative project. In: Henriksen K, Battles JB, Marks ES, Lewin DI, editors. Advances in Patient Safety: From Research to Implementation (Volume 2: Concepts and Methodology). Rockville, MD: Agency for Healthcare Research and Quality; 2005:255-278.

9. Scott D, Hallett C, Fettiplace R. Data-to-text summarisation of patient records: using computer-generated summaries to access patient histories. Patient Educ Couns 2013 Aug;92(2):153-159 [FREE Full text] [doi: 10.1016/j.pec.2013.04.019] [Medline: 23746770]

10. Armstrong S. The apps attempting to transfer NHS 111 online. Br Med J 2018 Jan 15;360:k156. [doi: 10.1136/bmj.k156] [Medline: 29335297]

11. Bachman JW. The patient-computer interview: a neglected tool that can aid the clinician. Mayo Clin Proc 2003 Jan;78(1):67-78. [doi: 10.4065/78.1.67] [Medline: 12528879]

12. Bingham P, Lilford RJ, Chard T. Strengths and weaknesses of direct patient interviewing by a microcomputer system in specialist gynaecological practice. Eur J Obstet Gynecol Reprod Biol 1984 Sep;18(1-2):43-56. [doi: 10.1016/0028-2243(84)90032-7] [Medline: 6548716]

13. Simmons Jr EM, Miller OW. Automated patient history-taking. Hospitals 1971 Nov 1;45(21):56-59. [Medline: 5095667]
14. Quaak MJ, Westerman RF, Schouten JA, Hasman A, van Bemmel JH. Computerization of the patient history—patient answers compared with medical records. Methods Inf Med 1986 Oct;25(4):222-228. [Medline: 3773779]

15. Schuman SH, Curry HB, Braunstein ML, Schneeweiss R, Jebaily GC, Glazer HM, et al. A computer-administered interview on life events: improving patient-doctor communication. J Fam Pract 1975 Aug;24(2):263-269. [Medline: 1185132]

16. Bachman J. Improving care with an automated patient history. Fam Pract Manag 2007;14(7):39-43 [FREE Full text] [Medline: 17696057]

17. Arora S, Goldberg AD, Menchine M. Patient impression and satisfaction of a self-administered, automated medical history-taking device in the emergency department. West J Emerg Med 2014 Feb;15(1):35-40 [FREE Full text] [doi: 10.1081/westjem.2013.2.11498] [Medline: 24695871]

18. Slack WW, Leviton A, Bennett SE, Fleischmann KH, Lawrence RS. Relation between age, education, and time to respond to questions in a computer-based medical interview. Comput Biomed Res 1988 Feb;21(1):78-84. [Medline: 3345654]

19. Greist JH, Gustafson DH, Stauss FF, Rowse GL, Laughren TP, Chiles JA. A computer interview for suicide-risk prediction. Am J Psychiatry 1973 Dec;130(12):1327-1332. [doi: 10.1176/ajp.130.12.1327] [Medline: 4585280]

20. Carr AC, Ghosh A, Ancill RJ. Can a computer take a psychiatric history? Psychol Med 1983 Feb;13(1):151-158. [doi: 10.1017/s003329170005157] [Medline: 6844461]

21. Paperny DM, Aono JY, Lehman RM, Hammar SL, Risser J. Computer-assisted detection and intervention in adolescent high-risk health behaviors. J Pediatr 1990 Mar;116(3):456-462. [doi: 10.1016/s0002-2476(05)82844-6] [Medline: 2308041]

22. Ammenwerth E, Schnell-Inderst P, Hoerbst A. Patient empowerment by electronic health records: first results of a systematic review on the benefit of patient portals. Stud Health Inform Technol 2011;165:63-67. [doi: 10.3233/978-1-60750-735-2-63] [Medline: 21685587]

23. Lancaster K, Abuozur A, Khaira M, Mathers A, Chan A, Bui V, et al. The use and effects of electronic health tools for patient self-monitoring and outcomes following release of medication use: systematic review. J Med Internet Res 2018 Dec 18;20(12):e294 [FREE Full text] [doi: 10.2196/ijmir.9284] [Medline: 30563822]

24. Semigran HL, Linder JA, Gidengil C, Mehrotra A. Evaluation of symptom checkers for self diagnosis and triage: audit study. Br Med J 2015 Jul 8;351:f3480 [FREE Full text] [doi: 10.1136/bmj.f3480] [Medline: 26157077]

25. Eysenbach G, CONSORT-EHEALTH Group. CONSORT-EHEALTH: improving and standardizing evaluation reports of web-based and mobile health interventions. J Med Internet Res 2011 Dec 31;13(4):e126 [FREE Full text] [doi: 10.2196/ijmir.1923] [Medline: 22209829]

26. DIANNA Diagnosis and Anamnesis Assistant. URL: http://www.diaana.fr [accessed 2019-08-09]

27. UpToDate. URL: http://www.uptodate.com [accessed 2019-07-22]

28. The National Center for Biotechnology Information. URL: https://www.ncbi.nlm.nih.gov/pubmed [accessed 2019-09-19]

29. Brukner P, Khan K. Brukner & Khan's Clinical Sports Medicine. Fourth Edition. New South Wales, Sydney: McGraw Hill Sports-Medicine; 2012.

30. Viechtbauer W, Smits L, Kotz D, Budé L, Spigt M, Serroyen J, et al. A simple formula for the calculation of sample size in pilot studies. J Clin Epidemiol 2015 Nov;68(11):1375-1379. [doi: 10.1016/j.jclinepi.2015.04.014] [Medline: 26146089]

31. Granja C, Janssen W, Johansen MA. Factors determining the success and failure of eHealth interventions: systematic review of the literature. J Med Internet Res 2018 May 1;20(5):e10235 [FREE Full text] [doi: 10.2196/jmir.2971] [Medline: 29716883]

32. Cahan A, Cimino JJ. A learning health care system using computer-aided diagnosis. J Med Internet Res 2017 Mar 8;19(3):e54 [FREE Full text] [doi: 10.2196/ijmir.6663] [Medline: 28274905]

33. Devarakonda MV, Mehta N, Tsou C, Liang JJ, Nowacki AS, Jelovsek JE. Automated problem list generation and physicians perspective from a pilot study. Int J Med Inform 2017 Sep;105:121-129 [FREE Full text] [doi: 10.1016/j.ijmedinf.2017.05.015] [Medline: 28750905]

34. Kostopoulos O, Rosen A, Round T, Wright E, Douiri A, Delaney B. Early diagnostic suggestions improve accuracy of GPs: a randomised controlled trial using computer-simulated patients. Br J Gen Pract 2015 Jan;65(630):e49-e54 [FREE Full text] [doi: 10.3399/bjgp15X196161] [Medline: 25548316]

35. Brodey B, Purcell SE, Rhea K, Maier P, First M, Zweede L, et al. Rapid and accurate behavioral health diagnostic screening: initial validation study of a web-based, self-report tool (the SAGE-SR). J Med Internet Res 2018 Mar;20(3):e108 [FREE Full text] [doi: 10.2196/ijmir.9428] [Medline: 29572204]

36. Ramnarayan P, Winrow A, Coren M, Nanduri V, Buchdahl R, Jacobs B, et al. Diagnostic omission errors in acute paediatric practice: impact of a reminder system on decision-making. BMC Med Inform Decis Mak 2006 Nov;6:37 [FREE Full text] [doi: 10.1186/1472-6947-6-37] [Medline: 17087835]

37. Moxpath C, Chambers N, Girling J, Garg S, Jelfs E, Brenner J. Perspectives on the enablers of e-heath adoption: an international interview study of leading practitioners. Health Serv Manag Res 2012 Aug;25(3):129-137. [doi: 10.1258/hsmr.2012.012018] [Medline: 23135887]

38. Meyer AN, Longhurst CA, Singh H. Crowdsourcing diagnosis for patients with undiagnosed illnesses: an evaluation of CrowdMed. J Med Internet Res 2016 Jan 14;18(1):e12 [FREE Full text] [doi: 10.2196/jmir.4887] [Medline: 26769236]

39. Graber ML, Tompkins D, Holland JJ. Resources medical students use to derive a differential diagnosis. Med Teach 2009 Jun;31(6):522-527. [doi: 10.1080/01421590802167436] [Medline: 19811168]
Abbreviations

**AMHTD**: automated medical history–taking device  
**DD**: differential diagnosis  
**DIAANA**: DIAgnosis & ANAmnesis  
**eHealth**: electronic health

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