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Adaptive e-learning environments for health professionals and students: a systematic review and meta-analysis

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learning environments for health professionals ts: a systematic review and meta-analysis

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33 TRANSPARENCY DECLARATION

The lead author (the manuscript's guarantor) affirms that the manuscript is an honest, accurate, and transparent account of the study being reported; that no important aspects of the study have been omitted; and that any discrepancies from the study as planned (and, if relevant, registered) have been explained.

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38 ABSTRACT (300 WORDS)

Objective: To systematically identify, appraise, and synthesize the evidence regarding the
effectiveness of adaptive e-learning environments (AEEs) in improving knowledge, competence,
and clinical behavior in health professionals and students.

Design: Systematic review and meta-analysis.

Methods: A search for studies published between January 2005 and April 2017 was performed in CINAHL, EMBASE, ERIC, PsycINFO, PubMed and Web of Science. Studies were eligible if they were controlled, and evaluated the effect of an AEE on knowledge, competence or clinical behavior in health professionals or students. AEEs were reviewed with regard to their clinical topic, theoretical framework, and adaptation process. Studies were included in the meta-analysis if they were randomized, and had a non-AEE control group. Effect sizes (ES) were pooled using a random effects model. Two authors screened studies, extracted data, assessed risk of bias, and coded quality of evidence independently.

Results: From a pool of 5,580 articles, we included 17 eligible studies enrolling 333 health professionals and 628 students. Risk of bias was generally high due to issues related to allocation concealment, similarity of baseline characteristics, blinding, and outcome data. Clinical topics were mostly related to diagnostic testing, theoretical frameworks were heterogeneous, and the adaptation process was characterized by 4 subdomains: goals, targeted variables, techniques, and timing. The pooled ES was 1.04 for knowledge (95% CI, -0.86-2.94; P.28), and 1.55 for competence (95% CI, 0.50-2.60; P.004). Statistical heterogeneity was high in all analyses. Conclusions: AEEs may improve only competence, not knowledge, in health professionals and

59 students. The adaptation process within AEEs may be more beneficial more learning

competencies since they are complex in nature, rather than learning knowledge, which generates

less cognitive load. Future research should report more clearly on the design and adaptation

process of AEEs, and target higher-level outcomes, such as clinical behavior.

PROSPERO registration number: CRD42017065585

Keywords: Computer-assisted instruction; medical education; nursing education; e-learning,

meta-analysis

66 ARTICLE SUMMARY

67	Strengths	and	Limitations	of	the	Study
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68	•	To our knowledge, this is the first systematic review and meta-analysis examining the
69		effectiveness of adaptive e-learning environments in improving knowledge, competence,
70		and clinical behavior in health professionals and students.

- Strengths of this review include the broad search strategy, and in-depth assessments of the
 risk of bias and the quality of evidence.
 - High statistical heterogeneity resulting from clinical and methodological diversity limits the interpretation of findings.
- Quantitative results should be treated with caution, given the small number and risk of
 bias of studies included in the meta-analysis.

77 INTRODUCTION

The use of information and communication technologies (ICTs) in the education of health professionals and students has become ubiquitous. Indeed, e-learning, defined as the use of ITCs to access educational curriculum and support learning¹, is increasingly present in clinical settings for the continuing education of health professionals²³, and in academic settings for the education of health professions students⁴. The interaction of health professionals and students with e-learning environments during the learning process generates a significant amount of data⁵. However, designers of e-learning environments and educators rarely make use of this data to optimize learning effectiveness and efficiency. Thus, in recent years, educational researchers have strived to develop e-learning environments that take a data-driven and personalized approach to education⁶⁻⁹. E-learning environments that take into account each user's interactions and performance level could anticipate what types of content and resources meet the user's needs, potentially increasing learning effectiveness and efficiency⁹. E-learning environments integrate information, in the form of text and multimedia (e.g., illustrations, animations, videos). They can include both asynchronous (i.e., designed for self-study) and synchronous (i.e., a class taught by an educator in real time) components¹. E-learning environments can be either *nonadaptive*, *adaptive*, or *intelligent*¹⁰ (see Figure 1). In the fields of computer science and educational technology, the term *adaptation* refers to the process executed

by a system based on ICTs of adapting educational curriculum content, structure or delivery to
the profile of a user ¹⁰.

97 Nonadaptive e-learning environments (Type A) provide a standardized training for all users.
98 While they can include instructional design variations (e.g., interactivity, feedback, practice

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exercises), they *do not* consider users' characteristics to provide a personalized training. They are generally considered to be as effective as non-e-learning educational interventions, such as largegroup classroom instruction and printed text, in improving learning outcomes^{11 12}. Adaptive e-learning environments (AEEs; **Type B1**) collect data to build each user's profile (e.g., navigation behavior, preferences, knowledge), and use simple techniques (e.g., adaptive information filtering, adaptive hypermedia) to adapt the content, navigation, presentation, multimedia, or strategies to provide personalized training^{7 8}. Intelligent e-learning environments (IEEs; **Type B2**) are a subtype of AEEs that use advanced adaptation techniques and user-modelling techniques derived from artificial intelligence (e.g., machine learning, rule-based systems, natural language processing) to provide a more personalized training for each user^{6 13-16}. In the context of this review, our use of the term "AEEs" includes IEEs. In recent years, AEEs have been developed and evaluated primarily in academic settings for students in mathematics, physics and related disciplines, for the acquisition of knowledge and development of cognitive skills (e.g., arithmetic calculation). Four meta-analyses reported on the effectiveness of AEEs among high school and university students in in these fields of study ¹³⁻¹⁵ ¹⁷. The results are promising: AEEs are in almost all cases more effective than large-group classroom instruction. In addition, Nesbit, et al.¹⁸ point out that AEEs are more effective than nonadaptive e-learning environments. The variability in the degree and the complexity of adaptation within AEEs mirrors the adaptation that can be observed in non–e-learning educational interventions (Type C). Some

interventions, like one-on-one human instruction and small-group classroom instruction,

generally have a high degree of adaptation since the instructor can adapt his teaching to the

individual profiles of learners and consider their feedback ¹⁹. Other interventions, like large-

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group classroom instruction, generally have a low degree of adaptation to individual leaners. In
some interventions, like paper-based instruction (e.g., handouts, textbooks), there is no adaptation
at all.
Despite evidence of the effectiveness of AEEs for knowledge acquisition and skill development

126 in areas such as mathematics in high school and university students, their effectiveness in

127 improving learning outcomes in health professionals and students has not yet been established.

128 To address this need, we conducted a systematic review and meta-analysis to identify and

quantitatively synthesize all comparative studies of AEEs involving health professionals andstudents.

131 Systematic Review and Meta-Analysis Objective

To systematically identify, appraise, and synthesize the best available evidence regarding the use
and the effectiveness of AEEs in improving knowledge, competence, and clinical behavior in
health professionals and students.

135 Systematic Review and Meta-Analysis Questions

136 We sought to answer the following questions with the systematic review:

1. What are the characteristics of studies assessing an AEE designed for health

138 professionals' and students' education?

139 2. What are the characteristics of AEEs designed for health professionals' or students'140 education?

141 We sought to answer the following question with the meta-analysis:

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2 3 4	142	3. What is the effectiveness of AEEs in improving knowledge, competence, and clinical
5 6	143	behavior in health professionals and students in comparison with nonadaptive e-learning
7 8	144	environments, and non-e-learning educational interventions?
9 10 11 12 12	145	METHODS
13 14 15	146	We planned and conducted this systematic review following the Effective Practice and
16 17	147	Organization of Care (EPOC) Cochrane Group guidelines ²⁰ , and reported it according to the
18 19 20	148	Preferred Reporting Items for Systematic review and Meta-Analysis (PRISMA) standards ²¹ (see
20 21 22	149	Supplementary File 1). We prospectively registered (International Prospective Register of
23 24	150	Systematic Reviews #CRD42017065585) and published the protocol of this systematic review ²²
25 26	151	²³ . Thus, in this paper, we present an abridged version of the methods with an emphasis on
27 28 29	152	changes made to the methods since the publication of the protocol.
30 31 32	153	Study Eligibility
33 34 35	154	We included primary research articles that assessed an AEE with licensed health professionals,
36 37	155	students, trainees, and residents in any discipline. We defined an AEE as a computer-based
38 39	156	learning environment which collects data to build each user's profile (e.g., navigation behavior,
40 41 42	157	individual objectives, knowledge), interprets these data through algorithms, and adapts in real-
43 44	158	time the content (e.g., showing/hiding information), navigation (e.g., specific links and paths),
45 46	159	presentation (e.g., page layout), multimedia presentation (e.g., images, videos), or tools (e.g.,
47 48 49	160	different set of strategies for different types of users) to provide a dynamic and evolutionary
50 51	161	learning path for each user ^{6 10} . We used the definitions of each type of adaptation proposed by
52 53	162	Knutov and colleagues ⁸ . We included AEEs with variable levels of technological complexity,
54 55 56 57 58	163	ranging from simple adaptive functionality (Type B1) to the use of artificial intelligence (IEEs,

Type B2). We considered for inclusion primary research articles in which the comparator was: 1)
165 a nonadaptive e-learning environment; 2) a non–e-learning educational intervention; 3) another
166 AEE with design variations. While included in the qualitative synthesis of the evidence for
167 descriptive purposes, the third comparator was excluded from the meta-analysis. Outcomes of
168 interest were knowledge, competence (including skills), and clinical behavior^{24 25}. Finally, in
169 terms of study design, we considered for inclusion all controlled, quantitative studies in
170 accordance with the EPOC Cochrane Review Group guidelines²⁶.

171 Study Identification

We previously published our search strategy²². Briefly, we designed a strategy in consultation with a librarian to search CINAHL, EMBASE, ERIC, PsycINFO, PubMed and Web of Science for primary research articles published between January 2005 and April 2017. We limited our search to articles published after 2005 since earlier studies seem to have a bias toward more positive results¹⁵, which could be explained by the effect of the novelty of e-learning on student motivation and learning outcomes. The search strategy revolved around 3 key concepts: "adaptive e-learning environments", "health professionals/students", and "effects on knowledge/competence/behavior" (see Supplementary file 2). To identify additional articles, we hand-searched 6 key journals (e.g., British Journal of Educational Technology, Computers and *Education*) and the reference lists of included primary research articles. We sought relevant articles published in English or French.

183 Study Selection

We worked independently and in duplicate (G.F. and M.-A.M.-C./T.M.) to screen all titles and
abstracts for inclusion using the EndNote software V8.0 (Clarivate Analytics). We resolved

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3 4	186	disagreements by consensus. We then performed the full-text assessment of potentially eligible
5 6 7	187	articles using the same methodology.
8 9 10	188	Data Extraction
11 12 13	189	One review author (G.F.) extracted data from included primary research articles using a modified
14 15	190	version of the data collection form developed by the EPOC Cochrane Review Group ²⁷ . The main
16 17	191	changes made to the extraction form were the addition of specific items relationg to the AEE
18 19	192	assessed in each study (e.g., adaptation techniques, duration of each training session). Two
20 21 22	193	review authors (T.M., MF.D.) validated the data extraction forms. For all studies, we extracted
23 24 25	194	the following data items if possible:
26 27	195	• the population and setting: study setting, study population, inclusion criteria, exclusion
28 29 30	196	criteria;
30 31 32	197	• <i>the methods</i> : study aim, study design, unit of allocation, study start date and end date, and
33 34 25	198	duration of participation;
35 36 37	199	• <i>the participants</i> : study sample, withdrawals and exclusions, age, sex, level of instruction,
38 39 40 41 42 43 44 45 46 47 48 49 50	200	number of years of experience as a health professional, practice setting, and previous
	201	experience using e-learning;
	202	• <i>the interventions</i> : name of intervention, theoretical framework, statistical model/algorithm
	203	used to generate the learning path, clinical topic, number of training sessions, duration of
	204	each training session, total duration of the training, adaptation techniques within the AEE
	205	(content, navigation, presentation, multimedia, tools), mode of delivery, presence of other
52 53	206	educational interventions and strategies;
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2 3 4	207	• the outcomes: name, time-points measured, definition, person measuring, unit of
5 6	208	measurement, scales, validation of measurement tool;
/ 8 9	209	• <i>the results</i> : results according to our primary (knowledge) and secondary (competence,
) 10 11	210	behavior) outcomes, comparison, time-point, baseline data, statistical methods used, and
12 13 14	211	key conclusions.
15 16	212	We contacted all of the corresponding authors (N=12) of the 17 included primary research
17 18 19	213	articles to validate the completed data extraction forms, and to provide us with missing data.
20 21 22	214	Assessment of the Risk of Bias
23 24 25	215	We worked independently and in duplicate (G.F. and T.M./MF.D.) to assess the risk of bias of
26 27	216	included primary research articles using the EPOC risk of bias criteria, based upon the data
28 29 30	217	extracted with the data collection form ²⁷ . A study was deemed at high risk of bias if the
30 31 32	218	individual criterion "random sequence generation" was scored at "high" or at "unclear" risk of
33 34 35	219	bias.
36 37 38	220	Data Synthesis
39 40	221	First, we synthesized data qualitatively using tables to provide an overview of the included
41 42 43 44	222	studies, and of the AEEs reported in these studies.
45 46	223	Second, using the Review Manager (RevMan) software V5.1, we first conducted a meta-analysis
47 48	224	to quantitatively synthesize the effectiveness of AEEs versus other educational interventions in
49 50	225	improving all learning outcomes. We included studies in the meta-analysis if the comparator
52 53	226	wasn't another AEE, if they were randomized, and if they reported outcome data. We then
54 55 56	227	conducted meta-analyses with the same comparison for each outcome for which data from at
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3 4	228	least 2 studies were available (i.e., knowledge, competence). For randomized controlled trials
5 6	229	(RCTs), we converted each post-test mean and standard deviation (SD) to a standardized mean
7 8 0	230	difference ([SMD], also known as Hedges g effect size [ES]). For crossover RCTs, we used
9 10 11	231	means pooled across each intervention. We pooled effect sizes using a random effects model.
12 13 14	232	Statistical significance was defined by a two-sided alpha of .05.
15 16 17	233	We first assessed heterogeneity qualitatively by examining the characteristics of included studies,
17 18 19	234	the similarities and disparities between the types of participants, the types of interventions, and
20 21	235	the types of outcomes. We then used the I^2 statistic within the RevMan software to quantify how
22 23	236	much the results varied across individual studies (i.e., between-study inconsistency, or
24 25 26	237	heterogeneity). We interpreted the I^2 values as follows: 0%-40%: might not be important; 30%-
27 28	238	60%: may represent moderate heterogeneity; 50%-90%: may represent substantial heterogeneity;
29 30	239	and 75%-100%: considerable heterogeneity ²⁸ . We performed sensitivity analysis to assess if the
31 32 22	240	exclusion of studies at high risk of bias or with a small sample size ($n<20$) would have had an
33 34 35	241	impact on statistical heterogeneity. The small number of studies included in each meta-analysis
36 37 38	242	did not allow for sensitivity of subgroup analyses to be performed.
39 40	243	Since less then 10 studies were included in the meta-analysis, we did not assess reporting biases
41 42 43	244	using a funnel plot, as suggested in the Cochrane Handbook ²⁹ .
44 45 46 47	245	Assessment of the Quality of Evidence
48 49	246	We worked independently and in duplicate (G.F. and MA.MC.) to assess the quality of
50 51	247	evidence for each individual outcome. We used the Grading of Recommendations Assessment,
52 53	248	Development, and Evaluation (GRADE) Web-based software, based upon the data extracted with
54 55 56 57	249	the data collection checklist ³⁰ . We considered 5 factors (risk of bias of included studies,
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250	indirectness of evidence, unexplained heterogeneity or inconsistency of results, imprecision of
251	the results, probability of reporting bias) for downgrading the quality of the body of evidence for
252	each outcome ³⁰ .
253	Patient and Public Involvement
254	Patients and the public were not involved in the selection of the research question and outcome
255	measures, or in the design of this systematic review and meta-analysis. No patients were involved
256	in the interpretation or writing up of results. We are unable to disseminate the results of the
257	research to study participants directly since this meta-analysis used aggregated data from
258	previous trials.
259	RESULTS
260	Study Flow
261	From a pool of 5,580 potentially relevant articles, we found 17 quantitative, controlled studies
262	assessing an AEE with health professionals or students (see Figure 2).
263	Out of 17 included studies in the qualitative synthesis, 4 studies compared two AEEs with design
264	variations ³¹⁻³⁴ , 4 studies were not randomized ³⁵⁻³⁸ , and 1 study had missing data ³⁹ . Thus, these 9
265	studies could not be included in the meta-analysis and the remaining 8 studies were used to
266	calculate an ES on learning outcomes.
267	Study Characteristics
268	We summarized the key characteristics of included studies in table format (see Table 1). In terms
269	of study population, in the 17 studies we have found published between 2006 and 2017,
270	investigators have evaluated AEEs mostly in the medical field, that is medical students $(n=7)^{35 37}$
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271	$^{40-43}$, medical residents (n=7) $^{31-34}$ 36 44 45 , nursing students (n=1) 46 , physicians in practice (n=1) 39 ,
272	and health sciences students $(n=1)^{47}$. The mean sample size was 56.53 participants (standard
273	deviation [SD] 44.13). In terms of study design, 13 out of 17 studies (77%) were randomized, 7
274	studies of which were randomized crossover trials ^{32 33 41-45} . The mean number of training
275	sessions was 2 (SD 1.07) and the mean training time was 4.33 hours (SD 3.79). Across the 17
276	studies, trainings with AEEs were spread over a mean period of time of 29.06 days (SD 44.91).
277	In terms of comparators, it is possible to underline three types of comparisons. The first
278	comparison is an AEE versus another AEE with design variations $(n=4)^{31-34}$, which implies that
279	one of the AEEs assessed had variations in its adaptation techniques (e.g., feedback in one AEE
280	is longer or more complex than in the other). The second comparison is an AEE versus a
281	nonadaptive e-learning environment (n=8) $35-374042-4447$. The third and final comparison is an
282	AEE versus another type of educational intervention, such as a paper-based educational
283	intervention, including handouts, textbooks or images $(n=3)^{394145}$, and a traditional educational
284	intervention, such as a group lecture $(n=1)^{46}$. As stated before, only the second and third types of
285	comparisons were included in the meta-analysis since our aim was to synthesize quantitatively th
286	effectiveness of AEEs versus other types of educational interventions. Finally, in terms of
287	outcomes, investigators evaluated learners' knowledge (n=11) ^{31 32 34 35 37 41-46} , satisfaction (n=9)
288	$^{31 33 35 37 41-45}$, competence (including skills) (n=6) $^{33 36 38-40 47}$, and metacognitive processes (n=3)
289	^{31 32 34} . However, none of the studies reported the assessment of health professionals' and
290	students' clinical behavior.

Table 1. Characteristics of included studies.

First author, year Country	Participants*	Study design [†]	No. and duration of training sessions	Duration of intervention	Comparison(s) [‡]	Outcome(s)§
Comparison: adaptive	e e-learning environm	nents vs. other educational interventions				
Cook, 2008 USA	R; N = 122	RXT; posttest-only, 4 groups	4 sessions; 30 minutes each	126 days	NEE	S, K
Crowley, 2010 USA	PP; N = 15	RCT; pretest-posttest, 2 groups	4 sessions; 4 hours each	138 days	Р	CΔ
Hayes-Roth, 2010 USA	MS, NS; N = 30	RCT; pretest-posttest-retention-test, 3 groups	NR; mean training time 2.36 hours	NR	1. NEE 2. NI	CΔ
Morente, 2013 Spain	NS; N = 73	RCT; pretest-posttest, 2 groups	1 session; 4 hours	1 day	L	KΔ
Munoz, 2010 Colombia	MS; N = 40	NRCT; pretest-posttest, 2 groups	NR; mean training time 5.97 hours	NR	NEE	S, K
Romito, 2016 USA	R; N = 24	NRCT; pretest-posttest-retention-test, 2 groups	1 session; 30 minutes	1 day	NEE & L	CΔ
Samulski, 2017 USA	MS, R, PP; N = 36	RXT; pretest-posttest, 2 groups	2 sessions; 20 minutes to 14 hours	30 days	Р	S, K
Thai, 2015 USA	HSC; N = 87	RCT; pretest-posttest-retention-test, 3 groups	1 session; 45 minutes	1 day	1. AEE 2. NEE	CΔ
Van Es, 2015 Australia	R; N = 43	RXT; posttest-only, 2 groups	3 sessions; NR	50 days	Р	S, K Δ
Van Es, 2016 Australia	MS; N = 46	RXT; posttest-only, 2 groups	3 sessions; 2 hours each	34 days	NEE	S, K
Wong, 2015 Australia	MS; N = 99	RXT; posttest-only, 2 groups	2 sessions; 1.5 hour each	14 days	NEE	S, KΔ
Wong, 2017 USA	MS; N = 178	NRCT; pretest-posttest-retention-test, 3 groups	1 session; NR	35 days	1. L 2. AEE & T	CΔ
Woo, 2006 USA	MS; N = 73	NRCT; pretest-posttest, 3 groups	1 session; 2 hours	1 day	1. NEE 2. NI	S, K
Comparison: adaptive	e e-learning vs. adapt	ive e-learning (two AEEs with design_variations)				
Crowley, 2007 USA	R; N = 21	RCT; pretest-posttest-retention-test, 2 groups	1 session; 4.5 hours	1 day	AEE	S, M, K
=						•••••••••••••••••••••••••••••••••••••••

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	El Saadawi, 2010 USA	R; N = 23	RXT; pretest–posttest, 2 groups	2 sessions; 2.25 hours each	2 days	AEE	M, K
	Feyzi-Begnagh, 2014 USA	R; N = 31	RCT; pretest–posttest, 2 groups	2 sessions; 2 & 3 hours	1 day	AEE	M, K
292 293 294 295 296 297 298 299	* Participants: MS indicate † Study design: RCT indic ‡ Comparison: AEE indica paper-based instruction (H § Outcomes: S indicates s improvement regarding th Included in the meta-and	es medical studen ates randomized o tes adaptive e-lea nandout, textbook, atisfaction; M, me is outcome in the alysis	ts; NS, nursing students; R, residents (physiciar controlled trial; RXT, randomized crossover trial arning environment; NEE, nonadaptive e-learnin , or latent image cases). tacognitive processes; K, knowledge; C, compe experimental group in comparison with the cont	ns in postgraduate training); PP, physicians ; NRCT, non-randomized controlled trial. g environment; NI, no-intervention control g tence (includes skills); B, behavior. Moreov rol group for studies comparing an AEE with	in practice; HSC, roup; L, large-grou er, Δ indicates a s h other educationa	health sciences str up classroom instru tatistically significa al interventions	udents. uction; P, ant
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300 Characteristics of Adaptive E-learning Environments

We summarized the key characteristics of AEEs assessed in the 17 studies in table format (see
 Table 2). In terms of the clinical topics of the AEEs, the majority of AEEs focused on training
 medical students and residents in executing and/or interpreting diagnostic tests. Indeed, the AEEs assessed focused on dermopathology and cytopathology microscopy ^{31-34 39 41 42 45} (n=8), on diagnostic imaging ^{36 43} (n=2), on chronic disease management ⁴⁴ (n=1), on pressure ulcer evaluation 46 (n=1), on the management of childhood illness 35 (n=1), on brief intervention for alcohol consumption ⁴⁰ (n=1), on electrocardiography ⁴⁷ (n=1), on fetal heart rate interpretation ³⁸ (n=1), and on hemodynamics 37 (n=1). Four out of 17 AEEs focused on developing the knowledge or skills of health professionals and students with regard to performing interventions in clinical practice ^{35 40 44 46}. Investigators adopted a wide variety of theoretical frameworks in the 17 studies. The most frequently used framework was cognitive tutoring, adopted in 5 studies ³¹⁻³⁴ ³⁹, which implies the use of a cognitive model. The integration of a cognitive model in an AEE implies the representation of all the knowledge in the field of interest in a way that is similar to the human mind for the purpose of understanding and predicting the cognitive processes of learners ⁴⁸. The second most used framework was perceptual learning, adopted in 3 studies ^{36 38 47}. Perceptual learning aims at improving information extraction skills of the environment and the development of automaticity in this respect in learners ³⁶. Theoretical frameworks relating to self-regulated learning ³⁴, learning styles ³⁵, guided mastery ⁴⁰, and cognitive load ⁴³ were also used. We propose 4 subdomains that emerged from the review to characterize the adaptation process of AEEs reported in the 17 studies: the goals of adaptation, the variables targeted by the adaptation process, the adaptation techniques, and the timing of the adaptation.

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5 4 5	322	First Subdomain: Goals of the Adaptation Process
5 6 7	323	This subdomain relates to the purpose of the adaptation process within the AEE. For most AEEs,
, 8 9	324	the adaptation process aims primarily to increase the effectiveness and/or efficiency of
10 11	325	knowledge acquisition and competence development relative to other training methods ^{31 34 35 37}
12 13	326	⁴⁰⁻⁴⁷ . For instance, several AEEs aimed to increase the diagnostic accuracy and reporting
14 15 16	327	performance of medical students and residents ^{31-33 36 38 39} . In cases where two adaptive AEEs
17 18	328	with certain variations in their techno-pedagogical design are compared with each other, the
19 20	329	adaptation process generally aims at improving the metacognitive and cognitive processes related
21 22 23	330	to learning ^{31 32 34} .
24 25 26	331	Second Subdomain: Variables Targeted by the Adaptation Process
27 28 20	332	This subdomain relates to the user-related data (variables) upon which the adaptation process is
29 30 31	333	based. The most frequently targeted variable is the user's scores after an assessment or a question
32 33	334	within the AEE (e.g., knowledge/skills scores, response accuracy scores) ^{35-38 40-47} . Other
34 35 26	335	frequently targeted variables include the user's actions during its use of the AEE (e.g., results of
36 37 38	336	problem-solving tasks, results of reporting tasks, requests for help) ^{31-34 39} , and the user's response
39 40 41	337	time regarding a specific question or task ^{36 38 47} .
42 43 44	338	Third Subdomain: Adaptation Techniques
45 46	339	The third subdomain relates to which adaptation techniques are mobilized in the AEE. In the
47 48	340	context of this review, the adaptation techniques are based upon the work of Knutov and
49 50 51	341	colleagues ⁸ . Content adaptation was the most used adaptation technique; it was implemented in
52 53	342	all AEEs reviewed (n=17). Content adaptation aims to adapt the textual information (curriculum
54 55 56 57	343	content) to the learner's profile through different mechanisms and to different degrees ⁸ .
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344 *Navigation adaptation* was the second most used adaptation technique (n=13). Navigation can be 345 adapted in two ways; it can be enforced or suggested. When enforced, an optimal personalized 346 learning path is determined for the user by the algorithms within the AEE. When suggested, there 347 are several personalized learning paths available to each user, who can determine the path he prefers himself⁸. Most reviewed studies included AEEs with enforced navigation, with one 348 349 optimal personalized learning path being determined through various algorithms. *Multimedia* 350 *adaptation* was the third most used adaptation technique (n=10). This adaptation technique, much 351 like content adaptation which relates to textual information, implies the adaptation of the 352 multimedia elements of the training such as videos, pictures, models, to the user's profile. 353 *Presentation adaptation* was the fourth most used adaptation technique (n=8). It implies the 354 adaptation of the layout of the page to the digital device used, or to the user's profile. *Tools* 355 *adaptation* was the least used adaptation technique (n=7). This technique results in providing a 356 different set of features or learning strategies for different types of users, such as different 357 interfaces for problem solving, and knowledge representation.

358 Fourth Subdomain: Timing of the Adaptation

This last subdomain relates to *when* the adaptation occurs during the learning process with the AEE. In 16 out of 17 studies, the adaptation occurred throughout the training with AEE, usually after an answer to a question or during intermediate problem-solving steps. In one study, adaptation techniques were only implemented at the beginning of the training with the AEE ³⁵.

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First author,				Adaptation proce	ess: four subdomains						
First author,											
year (Clinical Topic(s)	Theoretical framework(s)	Platform	First subdomain: Adaptation for	Second subdomain: Adaptation to which	Third Adap	suba statio	lomain n tech इ	: iniqu s	es	Fourth subdomain:
				which goals or objectives?	user-related variables?			LICOULIGUE			Timing of adaptation
Cook, 2008 h a	Diabetes, nyperlipidemia, asthma, depression	NR	NR	LEF	User knowledge	~	~				After each case-based question in each module (17 to 21 times/module).
Crowley, S 2007 d	Dermopathology; subepidermal vesicular dermatitis	Cognitive Tutoring	SlideTutor	LE, MC, DX.	User actions: results of problem-solving tasks; requests for help	1	1	1	1	1	At the beginning of each case
Crowley, D 2010 n	Dermopathology; melanoma	Cognitive Tutoring	SlideTutor	DX, RP.	User actions: results of problem-solving tasks; reporting tasks; requests for help	1	✓	1	1	1	At the beginning of each case
El Saadawi , E 2008 n	Dermopathology; melanoma	Cognitive Tutoring	ReportTutor	DX, RP.	User actions, report features	5	1	v	1		At the beginning of each case
El Saadawi, 2010	Dermopathology	Cognitive Tutoring	SlideTutor	DX, MC.	User actions: results of problem-solving tasks; reporting tasks; requests for help	1	1	1	1		During intermediate problem- solving steps.
Feyzi- E Begnagh, n 2014 d	Dermopathology; nodular and diffuse dermatitides	Cognitive Tutoring, Theories of Self- Regulated Learning	SlideTutor	LE, MC	User actions: results of problem-solving tasks; reporting tasks; requests for help	1	1	1	1	1	During each case or immediately after ech case.

Hayes-Roth, 2010	Brief intervention training in alcohol abuse	Guided Mastery	STAR Workshop	LE	User scores, user- generated dialogue	1	1				During clinical cases
Morente, 2013	Pressure ulcer evaluation	NR	ePULab	LE	User skills	~					Each pressure ulcer evaluation.
Munoz, 2010	Management of childhood illness	Learning Styles Framework	SIAS-ITS	LE, LEF	User knowledge, user learning style	1				1	At the beginning of the trainir
Romito, 2016	Transoesophageal echocardiography	Perceptual Learning	TOE PALM	DX	User response accuracy, user response time	1	1		1		After each clinical case.
Samulski, 2017	Cytopathology; pap test, squamous lesions, glandular lesions	NR	SmartSparrow	LE	User knowledge	1	1				During intermediate problem solving steps.
Thai, 2015	Electrocardiography	Perceptual Learing Theory; Adaptive response-time	PALM	LE, LEF	User response accuracy, user response time	1		1	1	1	After each user response.
Van Es, 2015	Diagnostic cytopathology; gynecology, fine needle aspiration, exfoliative fluid	NR	SmartSparrow	LE	User responses	V	1	1	1		During intermediate problem solving steps.
Van Es, 2016	Diagnostic cytopathology; gynecology, fine needle aspiration, exfoliative fluid	NR	SmartSparrow	LE	User responses	1	1	1	1	~	During intermediate problem solving steps.
Wong, 2015	Diagnostic imaging; chest X-rays, CT scans	Cognitive Load Theory	SmartSparrow	LE	User responses	1					During intermediate problem solving steps.

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	Wong, 2017	Fetal heart rate interpretation	Perceptual Learning	PALM	DX	User response accuracy, user response time	~	√	1	After each clinical case.
	Woo, 2006	Hemodynamics; baroreceptor reflex	NR	CIRCSIM-Tutor	LE	User knowledge, user responses	1	1	·	 After each user response
366 367 368	Goals/Objectiv NR indicates r	ves : LE indicates learni not reported.	ng effectiveness; LE	EF learning efficiency;	DX diagnostic	accuracy; MC metacognitive	gains; F	RP reporti	ng perform	ance.
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377 Risk of Bias Assessment

Results of included studies for the risk of bias assessment are presented in Figure 3 and Figure 4. In \geq 75% of studies, biases related to random sequence generation, similarity of baseline outcome measurements, and selective reporting of outcomes were low. Moreover, in \geq 50% of studies, biases related to the blinding of outcome assessment and contamination were low. Regarding the blinding of outcome assessment, in most studies, review authors judged that the outcome and the outcome measurement were not likely to be influenced by the lack of blinding, since studies had objective measures, i.e. an evaluative test of knowledge or competence. Regarding contamination bias, review authors scored studies at high risk if they had a crossover design.

However, in \geq 50% of studies, biases related to allocation concealment, similarity of baseline characteristics, blinding of participants and personnel, and incomplete outcome data were unclear or high. Review authors, by consensus, decided that the cut-off criterion in terms of risk of bias to be included in the meta-analysis was randomization of study participants. Thus, the four studies³⁵⁻³⁸ that presented high or unclear risk of bias regarding random sequence generation were not included in the meta-analysis.

40392Quantitative Results41

393 Effectiveness of AEEs versus other educational interventions in improving all learning outcomes

As we considered effect sizes larger than 0.8 to be large 49 , the pooled effect size (SMD 1.32;

395 95% confidence interval [CI] 0.20-2.43; Z =2.32, P 0.02) of AEEs compared to other educational

396 interventions in improving all learning outcomes suggests a significantly large effect (see Figure

5). However, significant statistical heterogeneity was observed among studies ($I^2 = 96\%$, P

398 <.00001), and individual effect sizes ranged from -1.10 to 3.05.

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2 3 4	399	Effectiveness of AEEs versus other educational interventions in improving knowledge
5 6 7	400	The pooled effect size (SMD 1.04; 95% CI -0.86-2.94; $Z = 1.08$, $P = 0.28$) of AEEs compared to
, 8 9	401	other educational interventions in improving knowledge suggests a large, but nonsignificant
10 11	402	effect (see Figure 6). Significant statistical heterogeneity was observed among studies (I^2 =98%, P
12 13 14	403	<.00001), and individual effect sizes ranged from -1.10 to 3.05. One study in particular ⁴⁴ reported
14 15 16	404	a negative effect size, but the difference between groups in knowledge scores was statistically
17 18	405	nonsignificant. Moreover, while participants using the AEE in the experimental group reported
19 20 21	406	the same knowledge scores than participants in the control group at the end of study, time spent
21 22 23	407	on instruction was reduced by 18% with the AEE compared to the nonadaptive e-learning
24 25 26	408	environment, thus improving learning efficiency ⁴⁴ .
27 28 29	409	Effectiveness of AEEs versus other educational interventions in improving competence
30 31	410	The pooled effect size (SMD 1.55; 95% CI 0.50-2.60; Z =2.90, P 0.004) of AEEs compared to
32 33	411	other educational interventions in improving competence suggests a significantly large effect (see
34 35 26	412	Figure 7). Statistical heterogeneity was lower than in previous analyses, but was still significant
36 37 38	413	($I^2 = 84\%$, P <.00001). Individual effect sizes ranged from 0.60 to 2.87.
40 41 42	414	Quality of the Evidence
43 44	415	The quality of evidence table produced with GRADE, as well as the justifications for each
45 46	416	decision, is presented in Supplementary File 3 (GRADE quality of evidence levels: very low, low,
47 48 40	417	moderate, high). For knowledge, the quality of evidence was deemed to be very low. More
49 50 51	418	precisely, risk of bias was deemed serious, inconsistency serious, indirectness not serious, and
52 53 54	419	imprecision serious. For competence, the quality of evidence was deemed to be low. More
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420 precisely, risk of bias was deemed serious, inconsistency serious, indirectness not serious, and421 imprecision serious.

DISCUSSION

423 Principal Findings

This is the first systematic review and meta-analysis to evaluate the effectiveness of AEEs in health professionals and students. We identified 17 relevant studies published since 2006, 13 of which assessed an AEE versus another educational intervention (large-group classroom instruction, nonadaptive e-learning environment or paper-based learning), and 4 of which assessed 2 AEEs with design variations head-to-head. When compared with other educational interventions, AEEs were associated with statistically significant improvements in learning outcomes in 9 out of 13 studies. Pooled ES were large for knowledge and competence, but only the latter was associated with a statistically significant effect. Statistical heterogeneity was high in all analyses. However, this finding is consistent with other meta-analyses in the field of medical education that also reported high heterogeneity across studies⁵⁰⁻⁵². A small number of eligible studies prohibited us from performing subgroup analyses which could have helped in explaining the source of the heterogeneity. The quality of evidence for all comparisons was either low or very low. Therefore, while we believe the results support the potential of AEEs for the education of health professionals and students, we recommend interpreting the ES with caution.

439 Comparison with Other Studies

440 To our knowledge, no previous systematic review and meta-analysis has specifically assessed the 441 effectiveness of AEEs in improving learning outcomes in health professionals and students, or Page 27 of 48

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any other population. However, interestingly, since the 1990's there has been a strong research
interest in the field of AEEs that integrate artificial intelligence (also known as intelligent elearning environments [IEEs] and intelligent tutoring systems [ITSs], **Type B2**) into elementary,
high school and postsecondary education for multiple subjects.¹⁵. Thus, multiple meta-analyses
have been conducted with regard to AEEs in that setting.

447 Steenbergen-Hu and Cooper ¹³ reported a mean ES of 0.35 of AEEs integrating artificial
448 intelligence on learning outcomes in college students when compared to all other types of
educational interventions. The mean ES was 0.37 when the comparator was large-group
classroom instruction, 0.35 when the comparator was nonadaptive e-learning, and 0.47 when the
451 comparator was textbooks or workbooks¹³.

Ma, et al. ¹⁶ reported a mean ES of 0.42 of AEEs integrating artificial intelligence on learning outcomes in elementary, high school and postsecondary students when compared to large-group classroom instruction. The mean ES was 0.57 when the comparator was nonadaptive e-learning, and 0.35 when the comparator was textbooks or workbooks. Interestingly, the mean ES was higher for studies which assessed an AEE in biology and physiology (0.59) and in humanities and social science (0.63) than in studies which assessed an AEE in mathematics (0.35) and physics (0.38)¹⁶.

Kulik and Fletcher ¹⁵ reported a mean ES of 0.65 of AEEs integrating artificial intelligence on
learning outcomes in elementary, high school, and postsecondary students when compared to
large-group classroom instruction. Education areas in this review were diverse (e.g.,
mathematics, computer science, physics), but none were related to health sciences. Interestingly,
the mean ES was 0.78 for studies up to 80 participants, and 0.30 for studies with more than 250

participants. Moreover, the mean ES for studies conducted with elementary and high school
 students was 0.44, compared to 0.75 for studies conducted with postsecondary students¹⁵.

Thus, in light of the results of these meta-analyses, the ES reported in our review may appear high. However, our review looked more specifically into the effectiveness of AEEs in improving learning outcomes in health professionals and students. This is significant since, in the metaanalyses of Steenbergen-Hu and Cooper¹³, Ma, et al.¹⁶, and Kulik and Fletcher¹⁵, AEEs seem to be more effective in postsecondary students ^{15 16} and for learning subjects related to biology, physiology and social science ¹⁶. Moreover, previous meta-analyses focused on the effectiveness of AEEs in improving procedural and declarative knowledge, and did not report on the effectiveness on AEEs in improving skills or competence. This is important since AEEs may be more effective for providing tailored guidance and coaching for developing skills and competence regarding complex clinical interventions, rather than learning factual knowledge, as the results of our review suggest.

477 Strengths and Limitations

Strengths of this systematic review and meta-analysis include the prospective registration and publication of a protocol based on rigorous methods in accordance with Cochrane and PRISMA guidelines; the exhaustive search in all relevant databases; the independent screening of the titles, abstracts and full-text of studies; the assessment of each included studies' risk of bias using EPOC Cochrane guidelines; and the assessment of the quality of evidence for each individual outcome using the GRADE methodology.

484 Our review also has limitations to consider. First, outcome measures varied widely across
485 studies. To address this issue, we conducted the meta-analysis using the SMD. Using the SMD

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allowed us to standardize the results of studies to a uniform scale before pooling them. However,
the SMD also has drawbacks, since this method assumes that the differences in SDs reflect
differences in outcome measures, and not differences attributable to variability among study
populations²⁹. Nevertheless, review authors judged that using the SMD was the best option for
this review, as it is the current practice in the field of knowledge synthesis in medical education¹²
⁵⁰.

492 Second, there was high inconsistency among study results, which we can mostly attribute to 493 differences in populations, AEE design, research methods, and outcomes. This resulted in 494 sometimes widely differing estimates of effect. To partly address this issue, we used a random-495 effects model for the meta-analysis, which assumes that the effects estimated in the studies are 496 different and follow a distribution²⁹. However, since a random-effects model awards more weight 497 to smaller studies to learn about the distribution of effects, it could potentially exacerbate the 498 effects of the bias in these studies²⁹.

Third, we made the decision to limit our search from the year 2005 onwards. This decision was based on the fact that studies published before 2005 seem to have a bias toward more positive results ¹⁵. This could be explained by the novelty of e-learning in earlier studies, which could have positively affected student motivation and learning outcomes. Moreover, educational technology has significantly evolved since the early 2000s.

Finally, publication bias could not be assessed by the means of a funnel plot since there were lessthan 10 studies included in the meta-analysis.

506 CONCLUSIONS

We found low and very low quality evidence that AEEs are associated with improved learning outcomes in health professionals and students in comparison with other educational interventions, such as nonadaptive e-learning environments and large-group classroom learning, across a range of topics. Heterogeneity was high across populations, interventions, comparators, outcomes and study designs.

In terms of *population*, future research should focus on assessing AEEs with health professionals in practice, such as registered nurses and physicians, rather than students in these disciplines. This could provide key insights into how AEEs can impact clinical behavior and, ultimately, patient outcomes. In terms of interventions, researchers should report more clearly on the goals of adaptation, the targeted variables by the adaptation process, the techniques of adaptation, and the timing of adaptation. Moreover, researchers should provide additional details regarding the underlining algorithms allowing the adaptation process in order to ensure replicability of findings. Regarding *comparators*, this review suggests there is a need for additional research using traditional comparators (i.e., large group classroom instruction) and more specific comparators (i.e., adaptive e-learning environment with design variations). Regarding outcomes and outcome measures, researchers should use validated measurement tools of knowledge. competence, and clinical behavior to facilitate knowledge synthesis. Moreover, the absence of studies assessing the impact of AEEs on health professionals' and students' clinical behavior demonstrates the need for further research with higher-level outcomes. Finally, in terms of *study designs*, researchers should focus on research designs allowing the assessment of the impact of multiple educational design variations and adaptation techniques within one study, such as factorial experiments.

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544 CONTRIBUTORSHIP STATEMENT

All review authors contributed to at least one aspect of each of the four criteria for authorshipdefined by the International Committee for Medical Journal Editors (ICJME).

547 G.F. contributed to the conception and design of the review, to the acquisition and analysis of

548 data, and to the interpretation of results. Moreover, G.F. drafted the initial manuscript. S.C.

549 contributed to the conception and design of the review, and to the interpretation of results. M.-

550 A.M.-C. contributed to the conception and design of the review, to the acquisition of data, and

interpretation of results. T.M. contributed to the conception and design of the review, to the acquisition of data, and to the interpretation of results, M.-F.D. contributed to the conception and design of the review, to the acquisition of data, and to the interpretation of results. G.M.-D. contributed to the conception and design of the review, and to the interpretation of results. J.C. contributed to the interpretation of results. M.-P.G. contributed to the interpretation of results. V.D. contributed to the interpretation of results. All review authors contributed to manuscript writing, critically revised the manuscript, gave final approval, and agreed to be accountable for all aspects of work, ensuring integrity and accuracy. **COMPETING INTERESTS** All authors have completed the ICMJE uniform disclosure form at www.icmje.org/coi disclosure.pdf and declare: no support from any organisation for the submitted work; no financial relationships with any organisation that might have an interest in the submitted work in the previous three years; no other relationships or activities that could appear to have influenced the submitted work. FUNDING This study received no funding. DATA SHARING STATEMENT No additional data are available.

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Figure 1. Types of educational interventions examined in the context of this review.

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Figure 3. Risk of bias summary: review authors' judgements about each risk of bias item for each included study.

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Figure 4. Risk of bias graph: review authors' judgements about each risk of bias item presented as percentages across all included studies.

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	Adapti	ve e-lear	ning	Other edu	icational i	nterv.	1	Std. Mean Difference	a Std. Mean Difference			
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Rando	m, 95% CI		
Cook 2008	76.2	0.9	62	77.2	0.9	62	13.1%	-1.10 [-1.48, -0.73]	+			
Hayes-Roth 2010	16.32	1.54	11	10.95	2.03	11	11.4%	2.87 [1.61, 4.12]				
Morente 2013	15.83	2.52	30	11.6	2.39	42	12.9%	1.71 [1.16, 2.26]		-		
Samulski 2017	5.83	2.43	18	4.47	2.47	18	12.7%	0.54 [-0.12, 1.21]		+- -		
Thai 2015	69	16	27	57	23	27	12.9%	0.60 [0.05, 1.14]		-		
Van Es 2015	91.8	2.3	12	82.6	4.3	15	11.9%	2.51 [1.46, 3.55]		_ _		
Van Es 2016	71.7	18.73	15	58.2	20.2	18	12.6%	0.67 [-0.03, 1.38]				
Wong 2015	68.2	2.7	39	60.5	2.3	41	12.7%	3.05 [2.39, 3.70]		-		
Total (95% CI)			214			234	100.0%	1.32 [0.20, 2.43]		•		
Heterogeneity: Tau ² =	2.42; Ch	i ² = 172	63, df =	7 (P < 0.0	0001); l ² -	96%			Lao L	<u> </u>		
Test for overall effect:	Z = 2.32	(P = 0.0)	2)						-10 -5 Other educational interv.	Adaptive e-learning	10	

Figure 5. Forest plot #1: meta-analysis of the effectiveness of adaptive e-learning environments versus other educational interventions in improving all learning outcomes.

242x56mm (300 x 300 DPI)

	Study or Subgroup	Adaptive e-	learning SD Total	Other edu Mean	icational int SD	erv. Total Weight	Std. Mean Difference IV, Random, 95% C	S	td. Mean Difference IV, Random, 95% CI	
	Cook 2008 Morente 2013	76.2 0 15.83 2.5).9 62 52 30	77.2 11.6	0.9 2.39	62 25.3% 42 25.0%	-1.10 [-1.48, -0.73 1.71 [1.16, 2.26		* +	
	Samulski 2017 Wong 2015	5.83 2.4 68.2 2	43 18 1.7 39	4.47 60.5	2.47 2.3	18 24.8% 41 24.8%	0.54 [-0.12, 1.21] 3.05 [2.39, 3.70]		* +	
	Total (95% CI)		149			163 100.0%	1.04 [-0.86, 2.94]	ĺ	-	
	Heterogeneity: Tau ² = Test for overall effect:	= 3.66; Chi ² = 1 :: Z = 1.08 (P =	145.70, df = 0.28)	= 3 (P < 0.0	0001); l ² = :	98%		-10 -5 Other education	al interv. Adaptive e-le	5 10
Figure (5. Forest plo	t #2: m	eta-a	nalysis	s of th	e effecti	iveness of a	daptive e-le	earning envi	ironments v
0		othe	er edu	ucatior	nal inte	erventio	ns in impro	ving knowle	edge.	
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	Adaptive e-learning					erv.	:	Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Hayes-Roth 2010	16.32	1.54	11	10.95	2.03	11	21.3%	2.87 [1.61, 4.12]	
Thai 2015	69	16	27	57	23	27	28.3%	0.60 [0.05, 1.14]	-
Van Es 2015	91.8	2.3	12	82.6	4.3	15	23.5%	2.51 [1.46, 3.55]	
Van Es 2016	71.7	18.73	15	58.2	20.2	18	26.9%	0.67 [-0.03, 1.38]	-
Total (95% CI)			65			71	100.0%	1.55 [0.50, 2.60]	◆
Heterogeneity. Tau ² = 0.93; Chi ² = 19.18, df = 3 (P = 0.0003); $I^2 = 84\%$									-10 -5 0 5 10
Test for overall effect:	Z = 2.90	(P = 0.0)	04)						Other educational interv. Adaptive e-learning

Figure 7. Forest plot #3: meta-analysis of the effectiveness of adaptive e-learning environments versus other educational interventions in improving competence.

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PRISMA 2009 Checklist

#	¥ (Checklist item	Reported on page #		
	1	Identify the report as a systematic review, meta-analysis, or both.	1		
mary 2	2 	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	4-5		
ON					
:	3 I	Describe the rationale for the review in the context of what is already known.	7-9		
4 Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).					
<u>.</u>	•				
gistration §	5 	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number.	10		
ı 6	6 S	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.	10-11		
rces	7 	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.	11		
6	3 I 1	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.	Supplementary File 2		
5	9 (i	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	11		
process 10	I C	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	11-12		
1'	1 I s	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	12		
ndividual 12	2 I	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	13		
ures 13	3	State the principal summary measures (e.g., risk ratio, difference in means).	13-14		
ults 14	4 I	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., I ²) for each meta-analysis.	13-14		
	Image: spin stration Image: spin stration Image: spin stration Ima	# 1 nary 2 DN 3 4 4 DN 3 4 4 Distration 5 1 6 rces 7 8 9 process 10 11 11 ndividual 12 sults 14	# Checklist item 1 Identify the report as a systematic review, meta-analysis, or both. nary 2 Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number. ON 3 Describe the rationale for the review in the context of what is already known. 4 Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS). jistration 5 Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number. i 6 Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale. cres 7 Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched. 9 State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis). 0rocesses for obtaining and confirming data frorm investigators. 11		

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PRISMA 2009 Checklist

3 4 5 Sec	ction/topic	#	Checklist item	Reported on page #
6 7 Risł 8	k of bias across studies	15	Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).	13
9 Add 10	litional analyses	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	14
1, RE	SULTS	<u>-</u>		
13 Stud 14	dy selection	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	15
15 Stud 16	dy characteristics	18	For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and provide the citations.	18
18 Risk	k of bias within studies	19	Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	27
¹⁹ Res 20	sults of individual studies	20	For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.	28-30
22 Syn	thesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency.	28-30
23 Risk	k of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).	28
25 Add 26	litional analysis	23	Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).	N/A
27 DIS	SCUSSION			
28 29 30	nmary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	30
31 Limi 32	itations	25	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of identified research, reporting bias).	
33 34 25	nclusions	26	Provide a general interpretation of the results in the context of other evidence, and implications for future research.	
36 FUI	NDING			
37 Fun 38	nding	27	Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the systematic review.	N/A
40 41 <i>From</i>	n: Moher D, Liberati A, Tetzlaff	J, Altm	an DG, The PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. PLoS	Med 6(7): e1000097.
42 doi:1	10.1371/journal.pmed1000097		For more information, visit: <u>www.prisma-statement.org</u> .	
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Supplementary	/ File 2 –	Search	Strategy	for	PubMed
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PubMed - 17 avril 2017

- ((Adaptive[TIAB] OR individualized [TIAB] OR personalized[TIAB] OR Tailored[TIAB]) AND (elearning[TIAB] OR learning[TIAB] OR Instruction[TIAB] OR "web-based instruction"[TIAB] OR computer-based instruction[TIAB] OR computer-based tutoring[TIAB] OR education[TIAB] OR tutorial[TIAB] OR tutorials[TIAB])) OR Intelligent tutoring system[TIAB] OR Intelligent tutoring systems[TIAB]
- 2. "Computer-Assisted Instruction"[MH]
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- 4. Health Personnel*[TIAB] OR Health professional*[TIAB] OR Health care profession*[TIAB] OR Healthcare profession*[TIAB] OR Medical student*[TIAB] OR Medical assistant*[TIAB] OR health worker*[TIAB] OR Audiologist*[TIAB] OR Chiropractor*[TIAB] OR Dentist[TIAB] OR Dentists[TIAB] OR Dietitian*[TIAB] OR Dermatolog*[TIAB] OR endocrinologist*[TIAB] OR Gastroenterolog*[TIAB]OR Gynecolog*[TIAB]OR Radiolog*[TIAB] OR Medical Staff[TIAB] OR Midwife*[TIAB] OR nutritionist*[TIAB] OR Nurse[TIAB] OR Nurses[TIAB] OR Optometrist*[TIAB] OR Occupational Therapist*[TIAB] OR Patholog*[TIAB] OR Paramedic[TIAB] OR Paediatric[TIAB] OR pediatrician*[TIAB] OR Paediatrician*[TIAB] OR pediatrist*[TIAB] OR pediatric[TIAB] OR Pharmacist*[TIAB] OR Pharmaconomist*[TIAB] OR Pharmacologist*[TIAB] OR Pharmacy technician*[TIAB] OR Phlebotomist*[TIAB] OR Physician OR Podiatrist*[TIAB] OR Psychologist*[TIAB] OR Psychotherapist*[TIAB] OR psychiatrist*[TIAB] OR Physical therapist*[TIAB] OR physiotherapist*[TIAB] OR Respiratory therapist*[TIAB] OR Surgeon*[TIAB] OR Clinician*[TIAB] OR Cardiologist*[TIAB] OR Emergency medical technician*[TIAB] OR emergency doctor*[TIAB] OR emergentologist*[TIAB] OR clinical officer*[TIAB] OR Community health worker*[TIAB] OR Radiographer*[TIAB] OR Surgical technologist*[TIAB] OR Radiotherapist*[TIAB] OR Anesthetist*[TIAB] OR Resident[TIAB] OR residents[TIAB]
- "Health Personnel"[MH] OR "Students, Premedical"[MH] OR "Students, Medical"[MH] OR "Students, Nursing"[Mesh]
- "Education, Premedical"[MH] OR "Education, Medical"[MH] OR "Education, Nursing"[MH] OR "Education, Pharmacy"[MH] OR "Education, Public Health Professional"[MH] OR "Clinical Clerkship"[MH]
- 7. #4 OR #5 OR 6
- knowledge*[TIAB] OR Aptitude*[TIAB] OR accuracy[TIAB] OR impact*[TIAB] OR skill*[TIAB] OR performance*[TIAB] OR Learning outcome*[TIAB] OR effectiveness[TIAB] OR efficacy[TIAB] OR improvement*[TIAB] OR Innovative*[TIAB] OR innovation*[TIAB] OR randomised controlled trial[TIAB] OR randomized controlled trial[TIAB]
- "Clinical Competence"[MH] "Quality Improvement"[MH] OR "Learning Curve"[MH] OR Knowledge [MH] OR "randomized controlled trial"[PT]
- 10. #8 OR 9
- 11. #3 AND #7 AND #10
- 12. (english[LA] OR french[LA]) AND 2005:2017[DP]
- 13. #11 AND #12

Résultats: 4 375

Supplementary File 3 – Summary of the quality of evidence

			Certainty	assessment			Nº of p	atients	Effect		Importance
№ of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Adaptive e- Learning Intervention	Other educational interventions	Absolute (95% Cl)	Certainty	
Knowledg	e										
4	randomised trials	serious ^{a,b}	serious ^c	not serious	serious ^d	none	149	163	SMD 1.04 SD higher (0.86 lower to 2.94 higher)	⊕○○○ VERY LOW	IMPORTANT
Competence										<u> </u>	
4	randomised trials	serious ^e	not serious	not serious	serious ^f	none	65	71	SMD 1.55 SD higher (0.5 higher to 2.6 higher)	⊕⊕⊖⊖ LOW	CRITICAL
CI: Confiden	ce interval; SMD): Standardised mea	n difference	I		80			I		I
Fynlan	ations										
	ations										
a. The four s	tudies were sco	red at unclear or high	n risk of bias regardin	g allocation concea	Iment. Thus, there is	s no guarantee that participants'	allocation to each grou	p didn't influence the d	elivery of the intervention.		
 The risk of the r	t bias for similari	ty of baseline measu	irements was uncleai	touch	s, groups in these st	tudies could be disproportionate	and the distribution may	y not be normal since s	sample size is generally small.		
d No author	mentioned in the	ese four studies that	the measurement in	touch.	dae was validated a	and no single instrument is used	multiple times. Accordi	on to sample size calcu	ulations, sample size was sufficier	nt in 3 studies	
e There was	a high risk of co	ontamination bias in	2 studies Incomplete	e outcome data for 2	potentially 3 studie	es. Participants that were more m	notivated that complete	d the studies may have	a induced a bias in the results		
f. One under	powered study.	according to authors	. and 2 studies for wh	hich the sample size	calculation is not m	nentioned. It is not mentioned in t	he measurement instru	ments were validated i	in the 4 studies, and no single ins	trument is used multiple times.	
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Efficacy of adaptive e-learning for health professionals and students: a systematic review and meta-analysis

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Date Submitted by the Author:	25-Mar-2019
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Primary Subject Heading :	Medical education and training
Secondary Subject Heading:	Health services research
Keywords:	MEDICAL EDUCATION & TRAINING, COMPUTER-ASSISTED INSTRUCTION, ADAPTIVE E-LEARNING ENVIRONMENTS, META- ANALYSIS, SYSTEMATIC REVIEW
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2 3 4	1	Efficacy of adaptive e-learning for health professionals and	
5 6 7	2	students: a systematic review and meta-analysis	
8 9 10	3		
11 12	4	Guillaume Fontaine ^{a, b*} , Sylvie Cossette ^{a, b} , Marc-André Maheu-Cadotte ^{a, b, c} ,	
13	5	Tanva Mailhot ^{a, b, d} , Marie-France Deschênes ^{a, e} , Gabrielle Mathieu-Dupuis ^f , José Côté ^{a, c}	>
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34 DECLARATION OF COMPETING INTERESTS

All authors have completed the ICMJE uniform disclosure form at www.icmje.org/coi_disclosure.pdf and declare: no support from any organisation for the submitted work; no financial relationships with any organisation that might have an interest in the submitted work in the previous three years; no other relationships or activities that could appear to have influenced the submitted work

40 TRANSPARENCY DECLARATION

The lead author (the manuscript's guarantor) affirms that the manuscript is an honest,
accurate, and transparent account of the study being reported; that no important aspects of the
study have been omitted; and that any discrepancies from the study as planned (and, if relevant,
registered) have been explained.

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2 3 4	45	FUNDING STATEMENT
5 6 7	46	This study received no funding.
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ABSTRACT (300 WORDS)

Objective: Although adaptive e-learning environments (AEEs) can provide personalized
instruction to health professional and students, their efficacy remains unclear. Therefore, this
review aimed to identify, appraise, and synthesize the evidence regarding the efficacy of AEEs in
improving knowledge, skills, and clinical behavior in health professionals and students.

53 **Design:** Systematic review and meta-analysis.

54 Data Sources: CINAHL, EMBASE, ERIC, PsycINFO, PubMed and Web of Science from the
55 first year of records to February 2019.

56 Eligibility Criteria: Controlled studies that evaluated the effect of an AEE on knowledge, skills
57 or clinical behavior in health professionals or students.

58 Screening, Data Extraction and Synthesis: AEEs were reviewed with regard to their topic, 59 theoretical framework, and adaptivity process. Studies were included in the meta-analysis if they 60 had a non-AEE control group and had no missing data. Effect sizes (ES) were pooled using a 61 random effects model. Two authors screened studies, extracted data, assessed risk of bias, and 62 coded quality of evidence independently.

Results: From a pool of 10,569 articles, we included 21 eligible studies enrolling 3,684 health
professionals and students. Clinical topics were mostly related to diagnostic testing, theoretical
frameworks were heterogeneous, and the adaptivity process was characterized by 5 subdomains:
method, goals, timing, factors, and types. The pooled ES was 0.70 for knowledge (95% CI, -0.081.49; *P*.08), and 1.19 for skills (95% CI, 0.59-1.79; *P* < .00001). Risk of bias was generally high.
Heterogeneity was large in all analyses.

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conclusions: AEEs appear particularly effective in improving skills in health professionals and udents. The adaptivity process within AEEs may be more beneficial for learning skills rather an factual knowledge, which generates less cognitive load. Future research should report more early on the design and adaptivity process of AEEs, and target higher-level outcomes, such as linical behavior.

ROSPERO registration number: CRD42017065585

. . . eywords: Computer-assisted instruction; medical education; nursing education; e-learning,

eta-analysis

77 ARTICLE SUMMARY

78 Strengths and Limitations of the Study

- This is the first systematic review and meta-analysis examining the efficacy of adaptive e learning environments in improving knowledge, skills, and clinical behavior in health
 professionals and students.
 - Strengths of this review include the broad search strategy, and in-depth assessments of the risk of bias and the quality of evidence.
 - High statistical heterogeneity resulting from clinical and methodological diversity limits the interpretation of findings.
- Quantitative results should be treated with caution, given the small number and risk of
 bias of studies included in the meta-analysis.

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88 INTRODUCTION

The use of information and communication technologies (ICTs) in the education of health professionals and students has become ubiquitous. Indeed, e-learning, defined as the use of ITCs to access educational curriculum and support learning¹, is increasingly present in clinical settings for the continuing education of health professionals^{2,3}, and in academic settings for the education of health professions students⁴. E-learning environments integrate information, in the form of text and multimedia (e.g., illustrations, animations, videos). They can include both asynchronous (i.e., designed for self-study) and synchronous (i.e., a class taught by an educator in real time) components¹. Nonadaptive e-learning environments, the most widespread type of e-learning environment today, provide a standardized training for all learners⁵⁶. While they can include instructional design variations (e.g., interactivity, feedback, practice exercises), they do not consider learners' characteristics and the data generated during the learning process to provide a personalized training⁶⁻⁸. This is problematic, since the interaction of health professionals and students with e-learning environments during the learning process generates a significant amount of data⁹. However, designers of e-learning environments and educators rarely make use of this data to optimize learning efficacy and efficiency⁹.

In recent years, educational researchers have strived to develop e-learning environments that take a data-driven and personalized approach to education¹⁰⁻¹³. E-learning environments that take into account each learner's interactions and performance level could anticipate what types of content and resources meet the learner's needs, potentially increasing learning efficacy and efficiency¹³. Adaptive e-learning environments (AEEs) were developed for this purpose. AEEs collect data to build each learner's profile (e.g., navigation behavior, preferences, knowledge), and use simple techniques (e.g., adaptive information filtering, adaptive hypermedia) to implement different

types of adaptivity targeting the content, navigation, presentation, multimedia, or strategies of the training to provide a personalized learning experience ¹¹¹². In the fields of computer science and educational technology, the term *adaptivity* refers to the process executed by a system based on ICTs of adapting educational curriculum content, structure or delivery to the profile of a learner¹⁴. Two main methods of adaptivity can be implemented within an AEE. The first method, *designed adaptivity*, is expert-based and refers to an educator who designs the optimal instructional sequence to guide learners to learning content mastery. The educator determines how the curriculum will adapt to learners based on a variety of factors, such as knowledge or response time to a question. This method of adaptivity is thus based on the expertise of the educator who specifies how technology will react in a particular situation on the basis of the "if THIS, then THAT" approach. The second method, *algorithmic adaptivity*, refers to use of algorithms to determine, for instance, the extent of the learner's knowledge and the optimal instructional sequence. Algorithmic adaptivity requires more advanced adaptivity techniques and learner-modelling techniques derived from the fields of computer science and artificial intelligence (e.g. Bayesian knowledge tracing, rule-based machine learning, natural language processing) 10 15-18.

127 The variability in the degree and the complexity of adaptivity within AEEs mirrors the adaptivity 128 that can be observed in non–e-learning educational interventions. Some interventions, like one-129 on-one human instruction and small-group classroom instruction, generally have a high degree of 130 adaptivity since the instructor can adapt his teaching to the individual profiles of learners and 131 consider their feedback ¹⁹. Other interventions, like large-group classroom instruction, generally 132 have a low degree of adaptivity to individual learners. In some interventions, like paper-based 133 instruction (e.g., handouts, textbooks), there is no adaptivity at all.

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AEEs have been developed and evaluated primarily in academic settings for students in mathematics, physics and related disciplines, for the acquisition of knowledge and development of cognitive skills (e.g., arithmetic calculation). Four meta-analyses reported on the efficacy of AEEs among high school and university students in in these fields of study ^{15-17 20}. The results are promising: AEEs are in almost all cases more effective than large-group classroom instruction. In addition, Nesbit, et al.²¹ point out that AEEs are more effective than nonadaptive e-learning environments. However, despite evidence of the efficacy of AEEs for knowledge acquisition and skill development in areas such as mathematics in high school and university students, their efficacy in improving learning outcomes in health professionals and students has not yet been established. To address this need, we conducted a systematic review and meta-analysis to identify and quantitatively synthesize all comparative studies of AEEs involving health professionals and students.

146 Systematic Review and Meta-Analysis Objective

To systematically identify, appraise, and synthesize the best available evidence regarding the
efficacy of AEEs in improving knowledge, skills, and clinical behavior in health professionals
and students.

150 Systematic Review and Meta-Analysis Questions

151 We sought to answer the following questions with the systematic review:

- 1. What are the characteristics of studies assessing an AEE designed for health professionals' and students' education?
 - 2. What are the characteristics of AEEs designed for health professionals' or students' education?

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156 We sought to answer the following question with the meta-analysis:

3. What is the efficacy of AEEs in improving knowledge, skills, and clinical behavior in
 health professionals and students in comparison with nonadaptive e-learning
 environments, and non–e-learning educational interventions?

METHODS

We planned and conducted this systematic review following the Effective Practice and
Organization of Care (EPOC) Cochrane Group guidelines²², and reported it according to the
Preferred Reporting Items for Systematic review and Meta-Analysis (PRISMA) standards²³ (see
Supplementary File 1). We prospectively registered (International Prospective Register of
Systematic Reviews #CRD42017065585) and published the protocol of this systematic review²⁴.
Thus, in this paper, we present an abridged version of the methods with an emphasis on changes
made to the methods since the publication of the protocol.

Study Eligibility

We included primary research articles that assessed an AEE with licensed health professionals, students, trainees, and residents in any discipline. We defined an AEE as a computer-based learning environment which collects data to build each learner's profile (e.g., navigation behavior, individual objectives, knowledge), interprets these data through algorithms, and adapts in real-time the content (e.g., showing/hiding information), navigation (e.g., specific links and paths), presentation (e.g., page layout), multimedia presentation (e.g., images, videos), or tools (e.g., different set of strategies for different types of learners) to provide a dynamic and evolutionary learning path for each learner^{10 14}. We used the definitions of each type of adaptivity proposed by Knutov and colleagues¹². We considered for inclusion studies in which AEEs had

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designed or algorithmic adaptivity, and studies including a co-intervention in addition to adaptive e-learning (e.g. paper-based instruction). We considered for inclusion primary research articles in which the comparator was: 1) a nonadaptive e-learning environment; 2) a non-e-learning educational intervention: 3) another AEE with design variations. While included in the qualitative synthesis of the evidence for descriptive purposes, the third comparator was excluded from the meta-analysis. Outcomes of interest were knowledge, skills, and behavior^{25 26}, and were defined as follows: 1) knowledge: subjective (e.g., learner self-report) or objective (e.g., multiple-choice question knowledge test) assessments of factual or conceptual understanding; 2) skills: subjective (eg, learner self-report) or objective (eg, faculty ratings) assessments of procedural skills (e.g. taking a blood sample, performing CPR) or cognitive skills (e.g. problem-solving, interpreting radiographs) in learners; 3) behavior: subjective (eg, learner self-report) or objective (eg, chart audit) assessments of behaviors in clinical practice (such as test ordering)⁶. In terms of study design, we considered for inclusion all controlled, quantitative studies in accordaSsnce with the EPOC Cochrane Review Group guidelines²⁷. We excluded studies that: 1) were not published in English or French; 2) were non-experimental; 3) were not controlled; 4) did not report on at least one of the outcomes of interest in this review; 5) did not have a topic related to the clinical aspects of health.

195 Study Identification

We previously published our search strategy²⁴. Briefly, we designed a strategy in consultation
with a librarian to search CINAHL, EMBASE, ERIC, PsycINFO, PubMed and Web of Science
for primary research articles published since the inception of each database up to February 2019.
The search strategy revolved around 3 key concepts: "adaptive e-learning environments", "health

200 professionals/students", and "effects on knowledge/competence (skills)/behavior" (see

201 Supplementary file 2). To identify additional articles, we hand-searched 6 key journals (e.g.,

202 British Journal of Educational Technology, Computers and Education) and the reference lists of

203 included primary research articles.

204 Study Selection

We worked independently and in duplicate (G.F. and M.-A.M.-C./T.M.) to screen all titles and abstracts for inclusion using the EndNote software V8.0 (Clarivate Analytics). We resolved disagreements by consensus. We then performed the full-text assessment of potentially eligible articles using the same methodology. Studies were included in the meta-analysis if they had a non-AEE control group and had no missing data.

210 Data Extraction

One review author (G.F.) extracted data from included primary research articles using a modified version of the data collection form developed by the EPOC Cochrane Review Group ²⁸. The main changes made to the extraction form were the addition of specific items relating to the AEE assessed in each study. Two review authors (T.M., M.-F.D.) validated the data extraction forms by reviewed the contents of each form against the data in the original article, adding comments when changes were needed. For all studies, we extracted the following data items if possible:

• *the population and setting*: study setting, study population, inclusion criteria, exclusion criteria;

• *the methods*: study aim, study design, unit of allocation, study start date and end date, and duration of participation;

1 2		
3 4	221	• <i>the participants</i> : study sample, withdrawals and exclusions, age, sex, level of instruction,
5 6	222	number of years of experience as a health professional, practice setting, and previous
7 8 0	223	experience using e-learning;
9 10 11	224	• <i>the interventions</i> : name of intervention, theoretical framework, statistical model/algorithm
12 13	225	used to generate the learning path, clinical topic, number of training sessions, duration of
14 15	226	each training session, total duration of the training, adaptivity subdomains (method, goals,
16 17 18	227	timing, factors, types), mode of delivery, presence of other educational interventions and
19 20	228	strategies;
21 22	229	• the outcomes: name, time-points measured, definition, person measuring, unit of
23 24 25	230	measurement, scales, validation of measurement tool;
26 27	231	• <i>the results</i> : results according to our primary (knowledge) and secondary (skills, behavior)
28 29 30 31 32	232	outcomes, comparison, time-point, baseline data, statistical methods used, and key
	233	conclusions.
33 34 35	234	We contacted the corresponding authors of included primary research articles to provide us with
36 37	235	missing data.
38 39 40 41	236	Assessment of the Risk of Bias
42 43	237	We worked independently and in duplicate (G.F. and T.M./MF.D.) to assess the risk of bias of
44 45 46	238	included primary research articles using the EPOC risk of bias criteria, based upon the data
47 48	239	extracted with the data collection form ²⁸ . A study was deemed at high risk of bias if the
49 50	240	individual criterion "random sequence generation" was scored at "high" or at "unclear" risk of
51 52 53	241	bias.
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242 Data Synthesis

First, we synthesized data qualitatively using tables to provide an overview of the includedstudies, and of the AEEs reported in these studies.

Second, using the Review Manager (RevMan) software V5.1, we first conducted a meta-analysis to quantitatively synthesize the efficacy of AEEs versus other educational interventions in improving all learning outcomes. We included studies in the meta-analysis if the comparator wasn't another AEE, if they were randomized, and if they reported outcome data. We then conducted meta-analyses with the same comparison for each outcome for which data from at least 2 studies were available (i.e., knowledge, skills). For randomized controlled trials (RCTs), we converted each post-test mean and standard deviation (SD) to a standardized mean difference ([SMD], also known as Hedges g effect size [ES]). For crossover RCTs, we used means pooled across each intervention. We pooled ESs using a random effects model. Statistical significance was defined by a two-sided alpha of .05.

We first assessed heterogeneity qualitatively by examining the characteristics of included studies, the similarities and disparities between the types of participants, the types of interventions, and the types of outcomes. We then used the l^2 statistic within the RevMan software to quantify how much the results varied across individual studies (i.e., between-study inconsistency, or heterogeneity). We interpreted the I² values as follows: 0%-40%: might not be important; 30%-60%: may represent moderate heterogeneity; 50%-90%: may represent substantial heterogeneity; and 75%-100%: considerable heterogeneity²⁹. We performed sensitivity analysis to assess if the exclusion of studies at high risk of bias or with a small sample size (n < 20) would have had an

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263 impact on statistical heterogeneity. The small number of studies included in each meta-analysis 264 did not allow for subgroup analyses to be performed.

265 Since less then 10 studies were included in the meta-analysis for each outcome, we did not assess 266 reporting biases using a funnel plot, as suggested in the Cochrane Handbook³⁰.

267 Assessment of the Quality of Evidence

We worked independently and in duplicate (G.F. and M.-A.M.-C.) to assess the quality of 268 269 evidence for each individual outcome. We used the Grading of Recommendations Assessment, 270 Development, and Evaluation (GRADE) Web-based software, based upon the data extracted with the data collection checklist ³¹. We considered 5 factors (risk of bias of included studies, 271 272 indirectness of evidence, unexplained heterogeneity or inconsistency of results, imprecision of 273 the results, probability of reporting bias) for downgrading the quality of the body of evidence for 1eg each outcome 31 . 274

275 **Patient and Public Involvement**

276 Patients and the public were not involved in setting the research question, the outcome measures, 277 the design or conduct of this systematic review. Patients and the public were not asked to advise 278 on interpretation of results or to contribute to the writing or editing of this document.

- RESULTS 279
 - 280 **Study Flow**

281 From a pool of 10,569 potentially relevant articles, we found 21 quantitative, controlled studies 282 assessing an AEE with health professionals or students (see Figure 1).

[Insert Figure 1]

Out of 21 included studies in the qualitative synthesis, 4 studies compared two AEEs with design variations³²⁻³⁵, and 4 studies had missing data³⁶⁻³⁹. Thus, these 8 studies could not be included in the meta-analysis and the remaining 13 studies were used to calculate an ES on learning outcomes.

288 Study Characteristics

We summarized the key characteristics of included studies in table format (see **Table 1**). In terms of study population, in the 21 studies examined published between 2003 and 2018, investigators have evaluated AEEs mostly in the medical field. Studies focused on medical students $(n=8)^{38-44}$, medical residents (n=8) 32-354145-47, physicians in practice (n=4) 36374148, nursing students (n=2) 4049 , nurses in practice (n=2) 4850 and health sciences students (n=1) 51 . Three studies focused on multiple populations ^{40 41 48}. The median sample size was 46 participants (interquartile range [IQR] 123). In terms of study design, 15 out of 21 studies (71%) were randomized, 7 studies of which were randomized crossover trials ^{33 34 41-43 45 47}. The median number of training sessions was 2 (IQR 2.5 sessions), the median training time was 2.13 hours (IQR 2.88 hours), and the median training period was 14 days (IQR 45 days). In terms of comparators, it is possible to underline three types of comparisons. The first comparison is an AEE versus another AEE with design variations $(n=4)^{32-35}$, which implies that one of the AEEs assessed had variations in its design, such as different types of adaptivity (e.g., feedback in one AEE is longer or more complex than in the other). The second comparison is an AEE versus a nonadaptive e-learning environment (n=11) ^{38-40 42-46 48 50 51}. The third and final comparison is an AEE versus another type of educational intervention, such as a paper-based educational intervention, including

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2 3 4	305	handouts, textbooks or images $(n=3)^{374147}$, or a traditional educational intervention, such as a
4 5 6	306	group lecture (n=2) 4952 . In one study, the comparator was not clearly reported 36 . As stated
7 8	307	before, only the second and third types of comparisons were included in the meta-analysis since
9 10 11	308	our aim was to synthesize quantitatively the efficacy of AEEs versus other types of educational
12 13	309	interventions. Finally, in terms of outcomes, investigators evaluated learners' knowledge (n=15)
14 15 16	310	³² ³³ ³⁵ ³⁶ ³⁸ ³⁹ ⁴¹⁻⁴⁵ ⁴⁷⁻⁵⁰ , satisfaction (n=9) ³² ³⁴ ³⁸ ³⁹ ⁴¹⁻⁴³ ⁴⁵ ⁴⁷ , skills (n=8) ³⁴ ³⁶ ³⁷ ⁴⁰ ⁴⁶ ⁵⁰⁻⁵² ,
16 17 18 19 21 22 24 25 27 28 29 31 32 33 45 36 7 38 30 41 23 44 45 46 78 90 51 22 34 55 57 58	311	netacognitive processes (n=3) ³² ³³ ³⁵ , attitudes (n=2) ³⁶ ⁵⁰ , and behavior (n=1) ⁵⁰ .
59 60		For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml

Table 1. Characteristics of included studies.

First author, year Country	Participants*	Study design [†]	No. and duration of training sessions	Duration of intervention	Comparison(s) [‡]	Outcome(s)§
Comparison: adapt	tive e-learning envi	ronments vs. other educational intervention	ns			
Casebeer, 2003 USA	PP; N = 181	RCT; posttest-only, 2 groups	4 sessions; 1 hour each	NR	NR	Knowledge Δ Attitudes Δ Skills Δ
Cook, 2008 USA	R; N = 122	RXT; posttest-only, 4 groups	4 sessions; 30 minutes each	126 days	NEE	Satisfaction Knowledge
Crowley, 2010 USA	PP; N = 15	RCT; pretest-posttest, 2 groups	4 sessions; 4 hours each	138 days	Ρ	Skills Δ
de Ruijter, 2018 Netherlands	NP; N = 269	RCT; pretest-posttest, 2 groups	No fixed sessions	180 days	NEE	Knowledge Attitudes Behavior Δ
Hayes-Roth, 2010 USA	MS, NS; N = 30	RCT; pretest-posttest-retention-test, 3 groups	NR; mean training time 2.36 hours	NR	1. NEE 2. NI	Skills Δ
Lee, 2017 USA	MS; N = 1522	NRCT; pretest-posttest, 3 groups	5 sessions; NR	42 days	NEE	Knowledge Skills Δ Behavior Δ
Micheel, 2017 USA	PP, NP; N = 751	NRCT; pretest–posttest–retention-test, 2 groups	NR	NR	NEE	Knowledge Δ
Morente, 2013 Spain	NS; N = 73	RCT; pretest–posttest, 2 groups	1 session; 4 hours	1 day	Т	Knowledge Δ
Munoz, 2010 Colombia	MS; N = 40	NRCT; pretest-posttest, 2 groups	NR; mean training time 5.97 hours	NR	NEE	Satisfaction Knowledge
Romito, 2016 USA	R; N = 24	NRCT; pretest–posttest–retention-test, 2 groups	1 session; 30 minutes	1 day	NEE & T	Skills Δ
Samulski, 2017 USA	MS, R, PP; N = 36	RXT; pretest–posttest, 2 groups	2 sessions; 20 minutes to 14 hours	1 month	Р	Satisfaction Knowledge
Thai, 2015 USA	HSC; N = 87	RCT; pretest–posttest–retention-test, 3 groups	1 session; 45 minutes	1 day	1. AEE 2. NEE	Skills Δ
Van Es, 2015 Australia	R; N = 43	RXT; posttest-only, 2 groups	3 sessions; NR	50 days	Р	Satisfaction Knowledge Δ
Van Es, 2016	MS; N = 46	RXT; posttest-only, 2 groups	3 sessions; 2 hours each	34 days	NEE	Satisfaction

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Wong, 2015 Australia MS; N = 99 RXT; postlest-only, 2 groups 2 sessions; 1.5 hour each 14 days NEE Satisfa Knowle Wong, 2017 USA MS; N = 178 NRCT; pretest-postlest-retention-test, 3 1 session; NR 35 days 1. T 2. AEE & T Skills / Skills / 2. AEE & T Woo, 2006 USA MS; N = 73 NRCT; pretest-postlest, 3 groups 1 session; 2 hours 1 day 1. NEE 2. NI Satisfa Knowle Comparison: adaptive e-learning vs. adaptive e-learning (two AEEs with design variations) 1 day AEE Satisfa Metacc proces Crowley, 2007 USA R; N = 21 RCT; pretest-postlest-retention-test, 2 groups 1 session; 4.5 hours 1 day AEE Satisfa Metacc proces El Saadawi, 2008 USA R; N = 20 RXT; pretest-postlest, 2 groups 2 sessions; 2 hours each 1 day AEE Satisfa Skills El Saadawi, 2010 USA R; N = 31 RCT; pretest-postlest, 2 groups 2 sessions; 2.25 hours each 2 days AEE Metacc process Knowle 2014 R; N = 31 RCT; pretest-postlest, 2 groups 2 sessions; 2.8 3 hours 1 day AEE Metacc process Knowle 313 * Participants: MS indicates medical students; NS, nursing students; R, residents (physicians in po	Wong, 2015 Australia MS; N = 99 RXT; postlest-only, 2 groups 2 sessions; 1.5 hour each 14 days NEE Satisfacting Knowledg Wong, 2017 USA MS; N = 178 NRCT; pretest-postlest-retention-test, 3 groups 1 session; NR 35 days 1. T 2. AEE & T Skills Δ Woo, 2006 USA MS; N = 73 NRCT; pretest-postlest, 3 groups 1 session; 2 hours 1 day 1. NEE Satisfacting 2. NI Comparison: adaptive e-learning vs. adaptive e-learning (vs. adaptive e-learning vs. adaptive e-learning (vs. AEE) The e-learning vs. adaptive e-learning vs. adaptive e-learning (vs. AEE) 1 session; 2 hours 1 day AEE Metacogn processes Knowledg USA R; N = 21 RCT; pretest-postlest, 2 groups 2 sessions; 2 hours each 1 day AEE Satisfacting Statisfacting Values USA R; N = 23 RXT; pretest-postlest, 2 groups 2 sessions; 2.25 hours each 1 day AEE Metacogn processes Knowledg 2014 R; N = 31 RCT; pretest-postlest, 2 groups 2 sessions; 2.83 hours 1 day AEE Metacogn processes Knowledg 3131 * Participants: MS indicates medical students; NS, nursing students; R, residents (physicians in postgraduate training); PP, physicians in practice; NP, nurses in practice; HP, nurses in practice; HP, inurses		Australia						Knowledge
Wong, 2017 USA MS; N = 178 NRCT; pretest-posttest-retention-test, 3 groups 1 session; NR 35 days 1. T 2. AEE & T Skills / Skills / 2. NI Woo, 2006 USA MS; N = 73 NRCT; pretest-posttest, 3 groups 1 session; 2 hours 1 day 1. NEE 2. NI Satisfa Knowle Comparison: adaptive e-learning vs. adaptive e-learning (two AEEs with design variations) I session; 4.5 hours 1 day AEE Metacc process Knowle Crowley, 2007 USA R; N = 21 RCT; pretest-posttest-retention-test, 2 groups 1 session; 4.5 hours 1 day AEE Satisfa Metacc process Knowle El Saadawi, 2008 USA R; N = 20 RXT; pretest-posttest, 2 groups 2 sessions; 2 hours each 1 day AEE Satisfa Metacc process Knowle El Saadawi, 2010 USA R; N = 23 RXT; pretest-posttest, 2 groups 2 sessions; 2.25 hours each 2 days AEE Metacc process Knowle 2014 R; N = 31 RCT; pretest-posttest, 2 groups 2 sessions; 2 & 3 hours 1 day AEE Metacc Process Knowle 1 * Participants: MS indicates medical students; NS, nursing students; R, residents (physicians in postgraduate training); PP, physicians in practice; NP, nurses in practice health sciences students. 1 day AEE Metacc Process Knowle <td>Wong, 2017 USA MS; N = 178 NRCT; pretest-posttest-retention-test, 3 groups 1 session; NR 35 days 1. T 2. AEE & T Skills Δ Woo, 2006 USA MS; N = 73 NRCT; pretest-posttest, 3 groups 1 session; NR 1 day 1. NEEE 2. NI Skills Δ Comparison: adaptive e-learning vs. adaptive e-learning (two AEEs with design variations) 1 day 2. NI Knowledg Crowley, 2007 USA R; N = 21 RCT; pretest-posttest-retention-test, 2 groups 1 session; 4.5 hours 1 day AEE Satisfactin Metacogn processee Knowledg El Saadawi, 2008 USA R; N = 20 RXT; pretest-posttest, 2 groups 2 sessions; 2 hours each 1 day AEE Satisfactin Metacogn processee Knowledg USA R; N = 23 RXT; pretest-posttest, 2 groups 2 sessions; 2.25 hours each 2 days AEE Metacogn processee Knowledg 2014 R; N = 31 RCT; pretest-posttest, 2 groups 2 sessions; 2 & 3 hours 1 day AEE Metacogn processee Knowledg 314 * Participants: MS indicates medical students; NS, nursing students; R, residents (physicians in postgraduate training); PP, physicians in practice; NP, nurses in practice; H heattris ciences students. * 1 day AEE Metacogn processee Knowledg * <td></td><td>Wong, 2015 Australia</td><td>MS; N = 99</td><td>RXT; posttest-only, 2 groups</td><td>2 sessions; 1.5 hour each</td><td>14 days</td><td>NEE</td><td>Satisfactio Knowledge</td></td>	Wong, 2017 USA MS; N = 178 NRCT; pretest-posttest-retention-test, 3 groups 1 session; NR 35 days 1. T 2. AEE & T Skills Δ Woo, 2006 USA MS; N = 73 NRCT; pretest-posttest, 3 groups 1 session; NR 1 day 1. NEEE 2. NI Skills Δ Comparison: adaptive e-learning vs. adaptive e-learning (two AEEs with design variations) 1 day 2. NI Knowledg Crowley, 2007 USA R; N = 21 RCT; pretest-posttest-retention-test, 2 groups 1 session; 4.5 hours 1 day AEE Satisfactin Metacogn processee Knowledg El Saadawi, 2008 USA R; N = 20 RXT; pretest-posttest, 2 groups 2 sessions; 2 hours each 1 day AEE Satisfactin Metacogn processee Knowledg USA R; N = 23 RXT; pretest-posttest, 2 groups 2 sessions; 2.25 hours each 2 days AEE Metacogn processee Knowledg 2014 R; N = 31 RCT; pretest-posttest, 2 groups 2 sessions; 2 & 3 hours 1 day AEE Metacogn processee Knowledg 314 * Participants: MS indicates medical students; NS, nursing students; R, residents (physicians in postgraduate training); PP, physicians in practice; NP, nurses in practice; H heattris ciences students. * 1 day AEE Metacogn processee Knowledg * <td></td> <td>Wong, 2015 Australia</td> <td>MS; N = 99</td> <td>RXT; posttest-only, 2 groups</td> <td>2 sessions; 1.5 hour each</td> <td>14 days</td> <td>NEE</td> <td>Satisfactio Knowledge</td>		Wong, 2015 Australia	MS; N = 99	RXT; posttest-only, 2 groups	2 sessions; 1.5 hour each	14 days	NEE	Satisfactio Knowledge
Woo, 2006 USA MS; N = 73 NRCT; pretest-posttest, 3 groups 1 session; 2 hours 1 day 1. NEE 2. NI Satisfa Comparison: adaptive e-learning vs. adaptive e-learning (two AEEs with design variations) Conversion: adaptive e-learning vs. adaptive e-learning (two AEEs with design variations) Crowley, 2007 USA R; N = 21 RCT; pretest-posttest-retention-test, 2 groups 1 session; 4.5 hours 1 day AEE Metacc process Knowle El Saadawi, 2008 USA R; N = 20 RXT; pretest-posttest, 2 groups 2 sessions; 2 hours each 1 day AEE Satisfa Metacc process Knowle El Saadawi, 2010 USA R; N = 23 RXT; pretest-posttest, 2 groups 2 sessions; 2.25 hours each 2 days AEE Metacc process Knowle Feyzi-Begnagh, 2014 R; N = 31 RCT; pretest-posttest, 2 groups 2 sessions; 2.8 3 hours 1 day AEE Metacc process Knowle 813 * Participants: MS indicates medical students; NS, nursing students; R, residents (physicians in postgraduate training); PP, physicians in practice; NP, nurses in practice health sciences students. * Participants: MS indicates randomized controlled trial; RXT, randomized crossover trial; NRCT, non-randomized controlled trial. * Comparison: AEE indicates randomized controlled trial; RXT, randomized crossover trial; NRCT, non-intervention control group; T, t	Woo, 2006 USA MS; N = 73 NRCT; pretest–posttest, 3 groups 1 session; 2 hours 1 day 1. NEE 2. NI Satisfactin Knowledg Comparison: adaptive e-learning vs. adaptive e-learning (two AEEs with design variations) Satisfactin Satisfactin Crowley, 2007 USA R; N = 21 RCT; pretest–posttest–retention-test, 2 groups 1 session; 4.5 hours 1 day AEE Satisfactin Metacogn processes Knowledg El Saadawi, 2008 USA R; N = 20 RXT; pretest–posttest, 2 groups 2 sessions; 2 hours each 1 day AEE Satisfactin Skills El Saadawi, 2010 USA R; N = 23 RXT; pretest–posttest, 2 groups 2 sessions; 2.25 hours each 2 days AEE Metacogn processes Knowledg Feyzi-Begnagh, 2014 Participants: MS indicates medical students; NS, nursing students; R, residents (physicians in postgraduate training); PP, physicians in practice; NP, nurses in practice; H health sciences students. Netacogn processes Knowledg 14 * Participants: MS indicates medical students; NS, nursing students; R, residents (physicians in postgraduate training); PP, physicians in practice; NP, nurses in practice; H health sciences students. 15 * Study design: RCT indicates aradomized controlled trial; RXT, randomized crossover trial; NRCT, non-randomized controlled trial. (comparison cher indicates adaptive e-learning environment; NLR, no-intervention control group; T, trad		Wong, 2017 USA	MS; N = 178	NRCT; pretest–posttest–retention-test, 3 groups	1 session; NR	35 days	1. T 2. AEE & T	Skills Δ
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 [§] Outcomes: ∆ indicates a statistically significant improvement regarding this outcome in the experimental group in comparison with the control group for studies comparison [§] AEE with other educational interventions. 	321	313 314 315	 Feyzi-Begnagh, 2014 USA * Participants: MS in bealth sciences str 	R; N = 31	RCT; pretest–posttest, 2 groups udents; NS, nursing students; R, residents (phy	2 sessions; 2 & 3 hours sicians in postgraduate training); PF	1 day P, physicians in p	AEE practice; NP, nurse	processes Knowledge es in practice; H
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322	Characteristics	of Adaptive	E-Learning	Environments
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We summarized the key characteristics of AEEs assessed in the 21 studies in table format (see
 Table 2). In terms of the clinical topics of the AEEs, the majority of AEEs focused on training
 medical students and residents in executing and/or interpreting diagnostic tests. Indeed, a significant proportion of the AEEs assessed focused on dermopathology and cytopathology microscopy ^{32-35 37 41 42 47} (n=8). Other topics were diagnostic imaging ^{43 46} (n=2), behavior change counseling ^{40 50} (n=2), chronic disease management ^{45 48} (n=2), pressure ulcer evaluation ⁴⁹ (n=1), childhood illness management 38 (n=1), electrocardiography 51 (n=1), fetal heart rate interpretation ⁵² (n=1), hemodynamics ³⁹ (n=1), chlamydia screening (n=1) ³⁶ and atrial fibrillation management $(n=1)^{44}$. The 21 AEEs examined were based on a wide variety of theoretical frameworks. The most frequently used framework was cognitive tutoring, adopted in 5 studies ^{32-35 37}, which refers to the use of a cognitive model. The integration of a cognitive model in an AEE implies the representation of all the knowledge in the field of interest in a way that is similar to the human mind for the purpose of understanding and predicting the cognitive processes of learners ⁵³. The second most used framework was perceptual learning, adopted in 3 studies ^{46 51 52}. Perceptual learning aims at improving information extraction skills of the environment and the development of automaticity in this respect in learners ⁴⁶. Interestingly, 2 studies used models from behavioral science, the Transtheoretical Model ³⁶ and the I-Change Model ⁵⁰, to tailor the AEE to the theoretical determinants of clinical behavior change in nurses and physicians in practice. Theoretical frameworks relating to self-regulated learning ³⁵, learning styles ^{38 48}, guided mastery ⁴⁰, and cognitive load ⁴³, problem-based-learning ³⁶, and situated learning ³⁶ were also used.

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344	Three main adaptive e-learning platforms were used by investigators in studies examined:
345	SlideTutor (n=4) ^{33 37 54 55} , Smart Sparrow (n=4) ^{41-43 47} , and the Perceptual Adaptive Learning
346	Module (PALM, n=3) ^{51 52 56} . SlideTutor is an AEE with algorithmic adaptivity which provides
347	cases to be solved by learners under supervision by the system. These cases incorporate
348	dermopathology virtual slides that must be examined by learners to formulate a diagnosis. An
349	expert knowledge base, consisting of evidence-diagnosis relationships, is used by SlideTutor to
350	create a dynamic solution graph representing the current state of the learning process and to
351	determine the optimal instructional sequence ⁵⁵ . Smart Sparrow is an AEE with designed
352	adaptivity which allows educators to determine adaptive factors, such as answers to questions,
353	response time to a question, and learner actions, to specify how the system will adapt the
354	instructional sequence or provide feedback. These custom learning paths can be more or less
355	personalized ⁴² . PALM is an AEE with algorithmic adaptivity aiming to improve perceptual
356	learning through adaptive response-time-based sequencing to determine dynamically the spacing
357	between different learning items based on each learner's accuracy and speed in interactive
358	learning trials ⁵¹ . Different custom adaptive e-learning platforms were used in other studies.
359	
Table 2. Characteristics of adaptive e-learning environments.

First		Theoretical		Adaptivity Subdomains					
Author, Year	Clinical Topic(s)	Framework(s)	Platform	Adaptivity Method	Adaptivity Goals	Adaptivity Timing	Adaptivity Factors	Adaptivity Types	
Casebeer, 2003	Chlamydia screening	Transtheoretical model of change; Problem-based learning; Situated learning theory	NR	Designed Adaptivity	To increase learning effectiveness (knowledge, skills).	Throughout the training, after case- based and practice- based questions.	User answers to questions	ContentNavigation	
Cook, 2008	Diabetes, hyperlipidemia, asthma, depression	NR	NR	Designed Adaptivity	To increase learning efficiency (knowledge gain divided by learning time).	After each case- based question in each module (17 to 21 times/module).	User knowledge	ContentNavigation	
Crowley, 2007	Dermopathology; subepidermal vesicular dermatitis	Cognitive Tutoring	SlideTutor	Algorithmic Adaptivity	To increase learning gains, metacognitive gains, and diagnostic performance.	At the beginning of each case.	User actions: results of problem- solving tasks; requests for help	 Content Navigation Presentation Multimedia Tools 	
Crowley, 2010	Dermopathology; melanoma	Cognitive Tutoring	SlideTutor	Algorithmic Adaptivity	To improve reporting performance and diagnostic accuracy.	At the beginning of each case.	User actions: results of problem- solving tasks; reporting tasks; requests for help	 Content Navigation Presentation Multimedia Tools 	
de Ruijter, 2018	Smoking cessation counseling	I-Change Model	Computer- Tailored E- Learning Program	Designed Adaptivity	To modify behavioral predictors and behavior.	At the beginning of the training.	Demographics, behavioral predictors, behavior	Content	
El Saadawi, 2008	Dermopathology; melanoma	Cognitive Tutoring	ReportTutor	Algorithmic Adaptivity	To teach how to correctly identify and document all relevant	At the beginning of each case.	User actions, report features	 Content Navigation Presentation Multimedia 	
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					prognostic factors in the diagnostic report.				
El Saadawi, 2010	Dermopathology	Cognitive Tutoring	SlideTutor	Algorithmic Adaptivity	To facilitate transfer of performance gains to real- world tasks that do not provide direct feedback on intermediate steps.	During intermediate problem-solving steps.	User actions: results of problem- solving tasks; reporting tasks; requests for help	• • •	Content Navigation Presentatio Multimedia
Feyzi- Begnagh, 2014	Dermopathology; nodular and diffuse dermatitis	Cognitive Tutoring, Theories of Self-Regulated Learning	SlideTutor	Algorithmic Adaptivity	To improve metacognitive and learning gains during problem solving.	During each case or immediately after each case.	User actions: results of problem- solving tasks; reporting tasks; requests for help	• • • • •	Content Navigation Presentatio Multimedia Tools
Hayes- Roth, 2010	Brief intervention training in alcohol abuse	Guided Mastery	STAR Workshop	NR	To improve attitudes and skills.	During clinical cases.	User scores, user- generated dialogue	•	Content Navigation
Lee, 2017	Treatment of atrial fibrillation	NR	Learning Assessment Platform	Designed Adaptivity	To increase learning effectiveness (knowledge, competence, confidence and practice).	After learning gaps identified in the first session.	Learning gaps in relation to objectives	•	Content
Micheel, 2017	Oncology	Learning Style Frameworks	Learning- style tailored educational platform	Designed Adaptivity	To increase learning effectiveness (knowledge).	After assessing the learning style.	Learning style	•	Presentatio Multimedia Tools

Morente, 2013	Pressure ulcer evaluation	NR	ePULab	Designed Adaptivity	To increase learning effectiveness (knowledge, skills).	Each pressure ulcer evaluation.	User skills	•	Content
Munoz, 2010	Management of childhood illness	Learning Styles Framework	SIAS-ITS	Designed Adaptivity	To increase learning effectiveness and efficiency.	At the beginning of the training.	User knowledge, user learning style	•	Content Tools
Romito, 2016	Transoesophageal echocardiography	Perceptual Learning	TOE PALM	Algorithmic Adaptivity	To improve response accuracy and response time.	After each clinical case.	User response accuracy, user response time	• •	Content Navigation Multimedia
Samulski, 2017	Cytopathology; pap test, squamous lesions, glandular lesions	NR	Smart Sparrow	Designed Adaptivity	To improve learning effectiveness.	During intermediate problem-solving steps.	User knowledge	•	Content Navigation
Thai, 2015	Electrocardiography	Perceptual Learing Theory; Adaptive response-time based algorithm	PALM	Algorithmic Adaptivity	To improve perceptual classification learning effectiveness and effiency.	After each user response.	User response accuracy, user response time	• • •	Content Presentatio Multimedia Tools
Van Es, 2015	Diagnostic cytopathology; gynecology, fine needle aspiration, exfoliative fluid	NR	Smart Sparrow	Designed Adaptivity	To improve learning effectiveness.	During intermediate problem-solving steps.	User responses	•	Content Navigation Presentatio Multimedia
Van Es, 2016	Diagnostic cytopathology; gynecology, fine needle aspiration, exfoliative fluid	NR	Smart Sparrow	Designed Adaptivity	To improve learning effectiveness.	During intermediate problem-solving steps.	User responses	• • •	Content Navigation Presentatio Multimedia Tools

	Wong, 2015	Diagnostic imaging; chest X-rays, CT scans	Cognitive Load Theory	Smart Sparrow	Designed Adaptivity	To improve learning effectiveness.	During intermediate problem-solving steps.	User responses	•	Content
) 1 2 3	Wong, 2017	Fetal heart rate interpretation	Perceptual Learning	PALM	Algorithmic Adaptivity	To improve response accuracy and response time.	After each clinical case.	User response accuracy, user response time	•	Content Navigation Multimedia
4 5 7 3	Woo, 2006	Hemodynamics; baroreceptor reflex	NR	CIRCSIM- Tutor	Algorithmic Adaptivity	To improve knowledge related to problem- solving tasks.	After each user response.	User knowledge, user responses	•	Content Navigation Tools
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We propose 5 subdomains that emerged from the review to characterize the adaptivity process of AEEs reported in the 21 studies: adaptivity method, adaptivity goals, adaptivity timing, adaptivity factors and adaptivity types.

368 First Subdomain: Adaptivity Method

This subdomain relates to the method of adaptivity that dictates how the AEE adapts instruction to a learner. As we previously described, there are two main methods of adaptivity: designed adaptivity and algorithmic adaptivity. The first is based on the expertise of the educator who specifies how technology will react in a particular situation on the basis of the "if THIS, then THAT" approach. The second refers to use of algorithms that will determine, for instance, the extent of the learner's knowledge and the optimal instructional sequence. In this review, 11 AEEs employed designed adaptivity ^{36 38 41-44 47-50 57}, and 9 AEEs employes algorithmic adaptivity ^{33 37 39} ^{51 52 54-56 58}. The adaptivity method wasn't specified in one study ⁴⁰.

3 377 Second Subdomain: Adaptivity Goals

This subdomain relates to the purpose of the adaptivity process within the AEE. For most AEEs, the adaptivity process aims primarily to increase the efficacy and/or efficiency of knowledge acquisition and skills development relative to other training methods ^{32 35 36 38-45 47-49 51}. For instance, several AEEs aimed to increase the diagnostic accuracy and reporting performance of medical students and residents ^{32-34 37 46 52}. In one study, the goal of adaptivity was to modify behavioral predictors and behavior in nurses ⁵⁰. In cases where two adaptive AEEs with certain variations in their techno-pedagogical design are compared with each other, the adaptivity process generally aims at improving the metacognitive and cognitive processes related to learning 32 33 35

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3 4	387	Third Subdomain: Adaptivity Timing
5 6 7	388	This subdomain relates to <i>when</i> the adaptivity occurs during the learning process with the AEE.
, 8 9	389	In 19 out of 21 studies, the adaptivity occurred throughout the training with AEE, usually after an
10 11	390	answer to a question or during intermediate problem-solving steps. However, in two studies,
12 13 14	391	adaptivity was only implemented at the beginning of the training with the AEE following survey
15 16 17	392	responses ^{38 50} .
18 19	393	Fourth Subdomain: Adaptivity Factors
20 21 22	394	This subdomain relates to the learner-related data (variables) upon which the adaptivity process is
23 24	395	based. The most frequently targeted variable is the learner's scores after an assessment or a
25 26 27	396	question within the AEE (e.g., knowledge/skills scores, response accuracy scores) ^{38-43 45-47 49 51 52} .
27 28 29	397	Other frequently targeted variables include the learner's actions during its use of the AEE (e.g.,
30 31	398	results of problem-solving tasks, results of reporting tasks, requests for help) ^{32-35 37} , and the
32 33 34	399	learner's response time regarding a specific question or task ^{46 51 52} .
35 36 37	400	Fifth Subdomain: Adaptivity Types
38 39	401	The final subdomain relates to which types of adaptivity are mobilized in the AEE: content,
40 41 42	402	navigation, multimedia, presentation and tools. In the context of this review, the adaptivity types
42 43 44	403	are based upon the work of Knutov and colleagues ¹² . Overall, 17 out of 21 (81%) AEEs
45 46	404	examined integrated more than one type of adaptivity. Content adaptivity was the most used
47 48 40	405	adaptivity type; it was implemented in all but one AEEs reviewed (n=20). Content adaptivity
49 50 51	406	aims to adapt the textual information (curriculum content) to the learner's profile through
52 53	407	different mechanisms and to different degrees ¹² . Navigation adaptivity was the second most used
54 55 56 57 58	408	adaptivity type (n=14). Navigation can be adapted in two ways; it can be enforced or suggested.

When enforced, an optimal personalized learning path is determined for the learner by an expert educator or by the algorithms within the AEE. When suggested, there are several personalized learning paths available to each learner, who can determine the path he prefers himself¹². Most reviewed studies included AEEs with enforced navigation, with one optimal personalized learning path being determined by an expert educator or by the algorithm. *Multimedia adaptivity* was the third most used adaptivity type (n=11). This adaptivity type, much like content adaptivity which relates to textual information, implies the adaptivity of the multimedia elements of the training such as videos, pictures, models, to the learner's profile. *Presentation adaptivity* was the fourth most used adaptivity type (n=9). It implies the adaptivity of the layout of the page to the digital device used, or to the learner's profile. *Tools adaptivity* was the least used adaptivity type (n=8). This technique results in providing a different set of features or learning strategies for different types of learners, such as different interfaces for problem solving, and knowledge 1.04 representation.

Risk of Bias Assessment

Results of included studies for the risk of bias assessment are presented in Figure 2 and Figure 3. In \geq 75% of studies, biases related to similarity of baseline outcome measurements, blinding of outcome assessment and selective reporting of outcomes were low. Moreover, in \geq 50% of studies, biases related to contamination were low. Regarding the blinding of outcome assessment, in most studies, review authors judged that the outcomes of interest and the outcome measurement were not likely to be influenced by the lack of blinding, since studies had objective measures, i.e. an evaluative test of knowledge or skills. Regarding contamination bias, review authors scored studies at high risk if they had a crossover design.

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2	40.1	
5 4	431	However, in \geq 50% of studies, biases related to random sequence generation, allocation
5 6	432	concealment, similarity of baseline characteristics, similarity of baseline characteristics, blinding
/ 8 0	433	of participants and personnel, and incomplete outcome data were unclear or high. Regarding
9 10 11	434	random sequence generation, an important number of studies did not report on the method of
12 13	435	randomization used by investigators. As per Cochrane recommendations, all eligibile studies
14 15	436	were incuded in the meta-analysis, regardless of the risk of bias assessment. Indeed, since almost
16 17	437	all studies scored overall at unclear risk of bias, Cochrane suggests to present an estimated
18 19 20	438	intervention effect based on all available studies, together with a description of the risk of bias in
21 22	439	individual domains ³⁰ .
23		
24 25	440	[Insert Figure 2]
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27 28	<i>11</i> 1	[Insert Figure 3]
29	441	[Insert Figure 5]
30 31	4.40	
32	442	Quantitative Results
33 34		
35	443	Efficacy of AEEs versus other educational interventions in improving knowledge
36		
37 38	444	The pooled ES (standardized mean difference [SMD] 0.70; 95% confidence interval [CI] -0.08-
39 40	445	1.49; Z =1.76, P 0.08) of AEEs compared to other educational interventions in improving
41 42	446	knowledge suggests a medium to large effect (see Figure 4). However, this result is not
43	4 4 77	
44 45	44 /	statistically significant. Significant statistical heterogeneity was observed among studies (I^2
46	448	=97%, P <.00001), and individual ESs ranged from -1.10 to 3.05. One study in particular 45
47 48		
49	449	reported a negative ES, but the difference between groups in knowledge scores was statistically
50 51	450	nonsignificant Moreover, while participants using the AFF in the experimental group reported
52	430	nonsignificant. Moreover, while participants using the ALE in the experimental group reported
53 54	451	the same knowledge scores as participants in the control group at the end of study, time spent on
55	450	instantion many hearth in 190/ with the AFF seminary data the many dentions a learning
56	452	instruction was reduced by 18% with the AEE compared to the honadaptive e-learning
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2 3 4	453	environment, thus improving learning efficiency ⁴⁵ . When that study ⁴⁵ is removed from the meta-
- 5 6	454	analysis, the pooled ES becomes statistically significant (SMD 1.07; 95% CI 0.28-1.85; Z =2.67,
7 8	455	<i>P</i> 0.008).
9 10 11 12	456	[Insert Figure 4]
13 14 15	457	Efficacy of AEEs versus other educational interventions in improving skills
16 17 18	458	As we considered ESs larger than 0.8 to be large ⁵⁹ , the pooled ES (SMD 1.19; 95% CI 0.59-
18 19 20	459	1.79; Z =3.88, P 0.0001) of AEEs compared to other educational interventions in improving
21 22	460	skills suggests a significantly large effect (see Figure 5). Statistical heterogeneity was lower than
23 24 25	461	in previous analyses, but was still significant (I^2 =89%, P <.00001). Individual ESs ranged from
25 26 27	462	0.17 to 2.87.
28 29	462	[Inset Figure 5]
30 31	403	[Insert Figure 5]
32 33 34	464	Quality of the Evidence
35 36	465	The quality of evidence table produced with GRADE, as well as the justifications for each
37 38 30	466	decision, is presented in Supplementary File 3 (GRADE quality of evidence levels: very low, low,
40 41	467	moderate, high). For knowledge, the quality of evidence was deemed to be very low. More
42 43	468	precisely, risk of bias was deemed serious, inconsistency serious, indirectness not serious, and
44 45	469	imprecision serious. For skills, the quality of evidence was deemed to be low. More precisely,
40 47 48	470	risk of bias was deemed serious, inconsistency serious, indirectness not serious, and imprecision
49 50	471	serious.
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DISCUSSION

473 Principal Findings

This is the first systematic review and meta-analysis to evaluate the efficacy of AEEs in health professionals and students. We identified 21 relevant studies published since 2003, 17 of which assessed an AEE versus another educational intervention (large-group classroom instruction, nonadaptive e-learning environment or paper-based learning), and 4 of which assessed 2 AEEs with design variations head-to-head. When compared with other educational interventions, AEEs were associated with statistically significant improvements in learning outcomes in 12 out of 17 studies. Pooled ESs were medium to large for knowledge and large for skills, but only the latter was associated with a statistically significant effect. Statistical heterogeneity was high in all analyses. However, this finding is consistent with other meta-analyses in the field of medical education that also reported high heterogeneity across studies ^{8 60 61}. A small number of eligible studies prohibited us from performing subgroup analyses which could have helped in explaining the source of the heterogeneity. The quality of evidence for all comparisons was either low or very low. Therefore, while we believe the results support the potential of AEEs for the education of health professionals and students, we recommend interpreting the ESs with caution.

Comparison with Other Studies

To our knowledge, no previous systematic review and meta-analysis has specifically assessed the efficacy of AEEs in improving learning outcomes in health professionals and students, or any other population. However, interestingly, since the 1990's there has been a strong research interest in the field of AEEs with algorithmic adaptivity (also known as intelligent learning environments [IEEs] or intelligent tutoring systems [ITSs]) into elementary, high school and

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494 postsecondary education for multiple subjects ¹⁷. Thus, multiple meta-analyses have been
495 conducted with regard to AEEs in that setting.

496 Steenbergen-Hu and Cooper ¹⁵ reported a mean ES of 0.35 of AEEs with algorithmic adaptivity
497 on learning outcomes in college students when compared to all other types of educational
498 interventions. The mean ES was 0.37 when the comparator was large-group classroom
499 instruction, 0.35 when the comparator was nonadaptive e-learning, and 0.47 when the comparator
500 was textbooks or workbooks¹⁵.

Ma, et al. ¹⁸ reported a mean ES of 0.42 of AEEs with algorithmic adaptivity on learning
outcomes in elementary, high school and postsecondary students when compared to large-group
classroom instruction. The mean ES was 0.57 when the comparator was nonadaptive e-learning,
and 0.35 when the comparator was textbooks or workbooks. Interestingly, the mean ES was
higher for studies which assessed an AEE in biology and physiology (0.59) and in humanities and
social science (0.63) than in studies which assessed an AEE in mathematics (0.35) and physics
(0.38)¹⁸.

Kulik and Fletcher ¹⁷ reported a mean ES of 0.65 of AEEs with algorithmic adaptivity on
learning outcomes in elementary, high school, and postsecondary students when compared to
large-group classroom instruction. Education areas in this review were diverse (e.g.,
mathematics, computer science, physics), but none were related to health sciences. Interestingly,
the mean ES was 0.78 for studies up to 80 participants, and 0.30 for studies with more than 250
participants. Moreover, the mean ES for studies conducted with elementary and high school
students was 0.44, compared to 0.75 for studies conducted with postsecondary students¹⁷.

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Thus, in light of the results of these meta-analyses, the ES reported in our review may appear high. However, our review looked more specifically into the efficacy of AEEs in improving learning outcomes in health professionals and students. This is significant since, in the metaanalyses of Steenbergen-Hu and Cooper¹⁵, Ma, et al.¹⁸, and Kulik and Fletcher¹⁷, AEEs seem to be more effective in postsecondary students ^{17 18} and for learning subjects related to biology, physiology and social science ¹⁸. Moreover, previous meta-analyses focused on the efficacy of AEEs in improving procedural and declarative knowledge, and did not report on the efficacy on AEEs in improving skills. This is important since AEEs may be more effective for providing tailored guidance and coaching for developing skills regarding complex clinical interventions, rather than learning factual knowledge, which generates less cognitive load ^{62 63}. Implications for Practice and Research This review provides important implications for the design and development of AEEs for health professionals and students. Table 3 presents 8 practical considerations for the design and

528 development of adaptive e-learning environments based on the results of this systematic review

529 for educators and educational researchers.

Table 3. Practical considerations for the design and development of adaptive e-learning environments.

Practical considerations	Explanations
Developing the Educational Content	• Given the adaptivity and the different learning pathways inherent to adaptive e- learning environments (AEEs), it is necessary to develop more pedagogical content (e.g. 60 minutes of learning) to reach the planned duration of each adaptive e-learning session (e.g. 30 minutes of learning).
Selecting a Theoretical Framework	• Selecting a theoretical framework coherent with the underlining principles of adaptivity of AEEs is crucial. These frameworks can be related to human cognition (e.g. Cognitive Load Theory, Cognitive Tutoring), behavior change (e.g. Transtheoretical model, I-Change Model) or learning (e.g. Perceptual Learning, Situated Learning).

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Selecting the Adaptivity Goal(s) • Selecting the adaptivity goal(s) is important, since it will dictate how the instruction be adapted in the AEE. The goal of adaptivity within an AEE may be to increase learning effectiveness, increase learning efficiency, modify behavioral predictors, of improve cognitive/metacognitive processes related to learning. Selecting the Adaptivity Timing • Selecting the timing of adaptivity within an AEE relates to when the adaptivity occu during the learning process. Adaptivity can be implemented at the beginning of the training only, or throughout the training. Adaptivity timing is closely linked to which adaptivity factor(s) Selecting the Adaptivity Factor(s) • Adaptivity factors are essentially data upon which the adaptivity process is based. These data can be related to the learner's performance (e.g. knowledge, skills), hi behavior / actions on the page (e.g. response time, requests for help), his overall learning path on the platform, or any other variables of interest in the learner. Selecting the Adaptivity Type(s) • Multiple types of adaptivity refers to the adaptation of the curiculum sequence; · Content adaptivity refers to the adaptation of the curiculum sequence; · Presentation adaptivity refers to the adaptation of alloy of the screen to the digital device used, or to the learner's profile; · Multimedia adaptivity refers to the adaptation of training features, learning strategi or learning assessment methods (e.g. interface for problem solving). Determining your technical resources and selecting the adaptive e-learning platform • After the content has been developed, the theoretical framework has been made, · cucial to determine your technical resources and equalute preexisting adaptive e- learning software to determine or ure tech	
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adaptivity process in order to ensure replicability of findings. Regarding *comparators*, this review suggests there is a need for additional research using traditional comparators (i.e., large group classroom instruction) and more specific comparators (i.e., adaptive e-learning environment with design variations). Regarding *outcomes* and outcome measures, researchers should use validated measurement tools of knowledge, skills, and clinical behavior to facilitate knowledge synthesis. Moreover, the very low number of studies assessing the impact of AEEs on health professionals' and students' clinical behavior demonstrates the need for further research with higher-level outcomes. Finally, in terms of *study designs*, researchers should focus on research designs allowing the assessment of the impact of multiple educational design variations and adaptivity types within one study, such as factorial experiments.

549 Strengths and Limitations

Strengths of this systematic review and meta-analysis include the prospective registration and publication of a protocol based on rigorous methods in accordance with Cochrane and PRISMA guidelines; the exhaustive search in all relevant databases; the independent screening of the titles, abstracts and full-text of studies; the assessment of each included studies' risk of bias using EPOC Cochrane guidelines; and the assessment of the quality of evidence for each individual outcome using the GRADE methodology.

556 Our review also has limitations to consider. First, outcome measures varied widely across 557 studies. To address this issue, we conducted the meta-analysis using the SMD. Using the SMD 558 allowed us to standardize the results of studies to a uniform scale before pooling them. Review 559 authors judged that using the SMD was the best option for this review, as it is the current practice 560 in the field of knowledge synthesis in medical education ⁶ ⁶⁰.

Second, there was high inconsistency among study results, which we can mostly attribute to differences in populations, AEE design, research methods, and outcomes. This resulted in sometimes widely differing estimates of effect. To partly address this issue, we used a randomeffects model for the meta-analysis, which assumes that the effects estimated in the studies are different and follow a distribution ³⁰. However, since a random-effects model awards more weight to smaller studies to learn about the distribution of effects, it could potentially exacerbate the effects of the bias in these studies ³⁰.

Finally, publication bias could not be assessed by the means of a funnel plot since there were lessthan 10 studies included in the meta-analysis.

570 CONCLUSIONS

Adaptive e-learning has significant potential to increase the effectiveness and efficiency of learning in health professionals and students. Through the different sub-domains of the adaptivity process (i.e. method, goals, timing, factors, types), AEEs can take into account the particularities inherent to each learner. This systematic review and meta-analysis underlines the potential of AEEs for improving knowledge and skills in health professionals and students in comparison with other educational interventions, such as nonadaptive e-learning environments and large-group classroom learning, across a range of topics. However, evidence was either of low or very low quality and heterogeneity was high across populations, interventions, comparators, and outcomes. Thus, additional comparative studies assessing the efficacy of AEEs in health professionals and students are needed to strengthen the quality of evidence.

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581 AUTHOR CONTRIBUTIONS

All review authors contributed to at least one aspect of each of the four criteria for authorship
defined by the International Committee for Medical Journal Editors (ICJME).

G.F. contributed to the conception and design of the review, to the acquisition and analysis of data, and to the interpretation of results. Moreover, G.F. drafted the initial manuscript, S.C. contributed to the conception and design of the review, and to the interpretation of results. M.-A.M.-C. contributed to the conception and design of the review, to the acquisition of data, and interpretation of results. T.M. contributed to the conception and design of the review, to the acquisition of data, and to the interpretation of results. M.-F.D. contributed to the conception and design of the review, to the acquisition of data, and to the interpretation of results, G.M.-D. contributed to the conception and design of the review, and to the interpretation of results. J.C. contributed to the interpretation of results. M.-P.G. contributed to the interpretation of results. V.D. contributed to the interpretation of results.

All review authors contributed to manuscript writing, critically revised the manuscript, gave final
approval, and agreed to be accountable for all aspects of work, ensuring integrity and accuracy.

596 DATA SHARING STATEMENT

597 No additional data available.

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616 Figure 3. Risk of bias graph: review authors' judgements about each risk of bias item presented as617 percentages across all included studies.

618 Figure 4. Forest plot representing the meta-analysis of the efficacy of adaptive e-learning versus 616 other educational interventions in improving knowledge.

620 Figure 5. Forest plot representing the meta-analysis of the efficacy of adaptive e-learning versus
 621 other educational interventions in improving skills.

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45	Figure 2. Risk of bias summary: review authors	ors'	jud	lger	nen	ts a	ιbοι	ut e	ach	risł	k of bias item for each included
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Figure 3. Risk of bias graph: review authors' judgements about each risk of bias item presented as percentages across all included studies.

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Adaptive e-learning				Other ed	ucational i	nterv.		Std. Mean Difference	Std. Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI	
Cook 2008	76.2	0.9	62	77.2	0.9	62	17.1%	-1.10 [-1.48, -0.73]		
de Ruijter 2018	14.876	1.242	121	14.8447	1.1093	103	17.4%	0.03 [-0.24, 0.29]	+	
Micheel 2017	7.861	1.691	282	7.461	1.691	317	17.6%	0.24 [0.08, 0.40]	•	
Morente 2013	15.83	2.52	30	11.6	2.39	42	16.3%	1.71 [1.16, 2.26]		
Samulski 2017	5.83	2.43	18	4.47	2.47	18	15.7%	0.54 [-0.12, 1.21]		
Wong 2015	68.2	2.7	39	60.5	2.3	41	15.8%	3.05 [2.39, 3.70]		
Total (95% CI)			552			583	100.0%	0.70 [-0.08, 1.49]	•	
Heterogeneity: Tau ² =	0.90; Chi	$^{2} = 150.$	19. df =	5 (P < 0.0)	$(0001); I^2 =$	97%		-		
Test for overall effect:	Z = 1.76	(P = 0.0)	8)						-4 -2 0 2 4	

Figure 4. Forest plot representing the meta-analysis of the efficacy of adaptive e-learning versus other educational interventions in improving knowledge.

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	Study or Subaroup	Adaptive e-learni Mean SD	ng O Total	ther educational Mean SD	interv. Total W	S eight	td. Mean Difference IV. Random, 95% CI	Std. Mean Difference IV. Random, 95% CI	
	Hayes-Roth 2010 Lee 2017	16.32 1.54 53 17	11 989	10.95 2.03 50 18	11 533	10.3%	2.87 [1.61, 4.12] 0.17 [0.07, 0.28]	•	
	Romito 2016 Thai 2015	67 4 69 16	12 27	53 6 57 23	12 27	11.1% 16.1%	2.65 [1.51, 3.80] 0.60 [0.05, 1.14]		
	Van Es 2015 Van Es 2016	91.8 2.3 71.7 18.73	12 15	82.6 4.3 58.2 20.2	15 18	11.9% 14.8%	2.51 [1.46, 3.55] 0.67 [-0.03, 1.38]		
	Wong 2017	0.55 0.22	39	0.46 0.21	86	17.3%	0.42 [0.04, 0.80]	-	
	Heterogeneity: Tau ² = Test for overall effect:	0.50; $Chi^2 = 57.00$ Z = 3.88 (P = 0.00	1105 , df = 6 (F 01)	P < 0.00001); I ² =	702 10 = 89%	00.0%	1.19 [0.59, 1.79]		
	restrict oreran eneet	2 - 5100 (1 - 6100						Other educational interv. Adaptive e-learning	
Figure 5	5. Forest plo	t represer	iting educa	the meta ational in	a-analy terven	sis o tion	of the efficac s in improvir	y of adaptive e-learning versus o g skills.	other
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PRISMA 2009 Checklist

Section/topic	#	Checklist item	Reported on page #
8 Title	1	Identify the report as a systematic review, meta-analysis, or both.	1
	-		
11 Structured summary 12 13	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	4-5
16 Rationale	3	Describe the rationale for the review in the context of what is already known.	7-9
17 Objectives 18	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).	9-10
20 METHODS	<u> </u>		
²¹ Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number.	10
24 Eligibility criteria 25	6	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.	10-11
²⁶ Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.	11-12
29 29 Search 30	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.	Supplementary File 2
3 Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	11-12
33 34 Data collection process 35	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	11-12
36 Data items 37	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	12-13
³⁸ Risk of bias in individual 10 studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	13
41 Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	13-14
42 43 44 44	14	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., I ²) for each meta-analysis.	13-14
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PRISMA 2009 Checklist

Section/topic	#	Checklist item	Reported on page #
Risk of bias across studies	15	Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).	13
Additional analyses	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	14
RESULTS	-		
3 Study selection	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	15
Study characteristics	18	For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and provide the citations.	18
8 Risk of bias within studies	19	Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	29-30
Results of individual studies	20	For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.	32-33
Synthesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency.	32-33
Risk of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).	31
Additional analysis	23	Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).	N/A
DISCUSSION			
Summary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	32-33
Limitations	25	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of identified research, reporting bias).	39
Conclusions	26	Provide a general interpretation of the results in the context of other evidence, and implications for future research.	36-39
Funding	27	Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the systematic review.	41
0 1 <i>From:</i> Moher D, Liberati A, Tetzlaff doi:10.1371/journal.pmed1000097	J, Altm	an DG, The PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. PLoS	Med 6(7): e1000097.

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Supplementary File 2 – Search Strategy for PubMed

PubMed – 17 avril 2017

- ((Adaptive[TIAB] OR individualized [TIAB] OR personalized[TIAB] OR Tailored[TIAB]) AND (elearning[TIAB] OR learning[TIAB] OR Instruction[TIAB] OR "web-based instruction"[TIAB] OR computer-based instruction[TIAB] OR computer-based tutoring[TIAB] OR education[TIAB] OR tutorial[TIAB] OR tutorials[TIAB])) OR Intelligent tutoring system[TIAB] OR Intelligent tutoring systems[TIAB]
- 2. "Computer-Assisted Instruction"[MH]
- 3. #1 OR #2
- Health Personnel*[TIAB] OR Health professional*[TIAB] OR Health care profession*[TIAB] OR 4. Healthcare profession*[TIAB] OR Medical student*[TIAB] OR Medical assistant*[TIAB] OR health worker*[TIAB] OR Audiologist*[TIAB] OR Chiropractor*[TIAB] OR Dentist[TIAB] OR Dentists[TIAB] OR Dietitian*[TIAB] OR Dermatolog*[TIAB] OR endocrinologist*[TIAB] OR Gastroenterolog*[TIAB]OR Gynecolog*[TIAB]OR Radiolog*[TIAB] OR Medical Staff[TIAB] OR Midwife*[TIAB] OR nutritionist*[TIAB] OR Nurse[TIAB] OR Nurses[TIAB] OR Optometrist*[TIAB] OR Occupational Therapist*[TIAB] OR Patholog*[TIAB] OR Paramedic[TIAB] OR Paediatric[TIAB] OR pediatrician*[TIAB] OR Paediatrician*[TIAB] OR pediatrist*[TIAB] OR pediatric[TIAB] OR Pharmacist*[TIAB] OR Pharmaconomist*[TIAB] OR Pharmacologist*[TIAB] OR Pharmacy technician*[TIAB] OR Phlebotomist*[TIAB] OR Physician OR Podiatrist*[TIAB] OR Psychologist*[TIAB] OR Psychotherapist*[TIAB] OR psychiatrist*[TIAB] OR Physical therapist*[TIAB] OR physiotherapist*[TIAB] OR Respiratory therapist*[TIAB] OR Surgeon*[TIAB] OR Clinician*[TIAB] OR Cardiologist*[TIAB] OR Emergency medical technician*[TIAB] OR emergency doctor*[TIAB] OR emergentologist*[TIAB] OR clinical officer*[TIAB] OR Community health worker*[TIAB] OR Radiographer*[TIAB] OR Surgical technologist*[TIAB] OR Radiotherapist*[TIAB] OR Anesthetist*[TIAB] OR Resident[TIAB] OR residents[TIAB]
- "Health Personnel"[MH] OR "Students, Premedical"[MH] OR "Students, Medical"[MH] OR "Students, Nursing"[Mesh]
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- knowledge*[TIAB] OR Aptitude*[TIAB] OR accuracy[TIAB] OR impact*[TIAB] OR skill*[TIAB] OR performance*[TIAB] OR Learning outcome*[TIAB] OR effectiveness[TIAB] OR efficacy[TIAB] OR improvement*[TIAB] OR Innovative*[TIAB] OR innovation*[TIAB] OR randomised controlled trial[TIAB] OR randomized controlled trial[TIAB]
- "Clinical Competence" [MH] "Quality Improvement" [MH] OR "Learning Curve" [MH] OR Knowledge [MH] OR "randomized controlled trial" [PT]
- 10. #8 OR 9
- 11. #3 AND #7 AND #10
- 12. (english[LA] OR french[LA]) AND 2005:2017[DP]
- 13. #11 AND #12

Résultats: 4 375

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Supplementary File 3 – Summary of the quality of evidence

Certainty assessment						№ of patients		Effect				
Nº of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	adaptive and intelligent e- learning environments	other educational interventions	Relative (95% Cl)	Absolute (95% Cl)	Certainty	Importance
Knowledge	Knowledge											
6	randomised trials	serious ^a	serious ^b	not serious	serious °	none	552	583	-	SMD 0.7 SD higher (0.08 lower to 1.49 higher)		IMPORTANT
Competence	e											
7	randomised trials	serious ^a	not serious	not serious	serious °	none	1105	702	-	SMD 1.19 SD higher (0.59 higher to 1.79 higher)		CRITICAL
CI: Confidence Explana a. Most studie may not be no b. Studies yiel c. Most studie	I: Confidence interval; SMD: Standardised mean difference Explanations I: Most studies have unclear or high risk of bias with regard to random sequence generation and allocation concealment. The risk of bias for similarity of baseline measurements was unclear for some studies. Thus, groups in these studies could be disproportionate and the distribution have not be normal since sample size is generally small. I: Studies yield widely differing estimates of effect (heterogeneity or variability in results). The individual confidence intervals of some studies almost do not touch. I: Most studies include few participants and few events and have wide confidence intervals. Measurement instruments often not validated. Sample size often unsufficient.											
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BMJ Open

Efficacy of adaptive e-learning for health professionals and students: a systematic review and meta-analysis

Journal:	BMJ Open
Manuscript ID	bmjopen-2018-025252.R2
Article Type:	Research
Date Submitted by the Author:	10-Jun-2019
Complete List of Authors:	Fontaine, Guillaume; University of Montreal, Faculty of Nursing; Montreal Heart Institute, Research Center Cossette, Sylvie ; University of Montreal, Faculty of Nursing; Montreal Heart Institute, Research Center Maheu-Cadotte, Marc-André; University of Montreal, Faculty of Nursing; Montreal Heart Institute, Research Center Mailhot, Tanya; Northeastern University, Bouve College of Health Sciences; Montreal Heart Institute, Research Center Deschênes, Marie-France; University of Montreal, Faculty of Nursing Mathieu-Dupuis, Gabrielle; University of Montreal, Faculty of Nursing Mathieu-Dupuis, Gabrielle; University of Montreal, School of Librarianship and Information Science Côté, José; University of Montreal, Faculty of Nursing; University of Montreal Hospital Centre, Research Center Gagnon, Marie-Pierre; Laval University, Faculty of Nursing; CHU de Québec-Université Laval, Research Center Dubé, Veronique; University of Montreal, Faculty of Nursing; University of Montreal Hospital Centre, Research Center
Primary Subject Heading :	Medical education and training
Secondary Subject Heading:	Health services research
Keywords:	MEDICAL EDUCATION & TRAINING, COMPUTER-ASSISTED INSTRUCTION, ADAPTIVE E-LEARNING ENVIRONMENTS, META- ANALYSIS, SYSTEMATIC REVIEW

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Page 1 of 54

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2 3 4	1	Efficacy of adaptive e-learning for health professionals and
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11 12	4	Guillaume Fontaine ^{a, b*} , Sylvie Cossette ^{a, b} , Marc-André Maheu-Cadotte ^{a, b, c} ,
13 14	5	Tanya Mailhot ^{a, b, d} , Marie-France Deschênes ^{a, e} , Gabrielle Mathieu-Dupuis ^f , José Côté ^{a, c} ,
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53 54 55 56 57 58	22	Word count (excluding title page, abstract, references, figures, and tables): 5,823 words
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ABSTRACT (300 WORDS)

49 Objective: Although adaptive e-learning environments (AEEs) can provide personalized
50 instruction to health professional and students, their efficacy remains unclear. Therefore, this
51 review aimed to identify, appraise, and synthesize the evidence regarding the efficacy of AEEs in
52 improving knowledge, skills, and clinical behavior in health professionals and students.

53 **Design:** Systematic review and meta-analysis.

54 Data Sources: CINAHL, EMBASE, ERIC, PsycINFO, PubMed and Web of Science from the
55 first year of records to February 2019.

56 Eligibility Criteria: Controlled studies that evaluated the effect of an AEE on knowledge, skills
57 or clinical behavior in health professionals or students.

Screening, Data Extraction and Synthesis: Two authors screened studies, extracted data,
assessed risk of bias, and coded quality of evidence independently. AEEs were reviewed with
regard to their topic, theoretical framework, and adaptivity process. Studies were included in the
meta-analysis if they had a non-AEE control group and had no missing data. Effect sizes (ES)
were pooled using a random effects model.

Results: From a pool of 10,569 articles, we included 21 eligible studies enrolling 3,684 health
professionals and students. Clinical topics were mostly related to diagnostic testing, theoretical
frameworks were varied, and the adaptivity process was characterized by 5 subdomains: method,
goals, timing, factors, and types. The pooled ES was 0.70 for knowledge (95% CI, -0.08-1.49; *P*.08), and 1.19 for skills (95% CI, 0.59-1.79; *P* < .00001). Risk of bias was generally high.

68 Heterogeneity was large in all analyses.

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onclusions: AEEs appear particularly effective in improving skills in health professionals and idents. The adaptivity process within AEEs may be more beneficial for learning skills rather an factual knowledge, which generates less cognitive load. Future research should report more early on the design and adaptivity process of AEEs, and target higher-level outcomes, such as nical behavior.

ROSPERO registration number: CRD42017065585

. . C. eywords: Computer-assisted instruction; medical education; nursing education; e-learning,

eta-analysis

77 ARTICLE SUMMARY

78 Strengths and Limitations of the Study

- This is the first systematic review and meta-analysis examining the efficacy of adaptive e learning environments in improving knowledge, skills, and clinical behavior in health
 professionals and students.
 - Strengths of this review include the broad search strategy, and in-depth assessments of the risk of bias and the quality of evidence.
 - High statistical heterogeneity resulting from clinical and methodological diversity limits the interpretation of findings.
- Quantitative results should be treated with caution, given the small number and risk of
 bias of studies included in the meta-analysis.

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88 INTRODUCTION

The use of information and communication technologies (ICTs) in the education of health professionals and students has become ubiquitous. Indeed, e-learning, defined as the use of ITCs to access educational curriculum and support learning¹, is increasingly present in clinical settings for the continuing education of health professionals^{2,3}, and in academic settings for the education of health professions students⁴. E-learning environments integrate information, in the form of text and multimedia (e.g., illustrations, animations, videos). They can include both asynchronous (i.e., designed for self-study) and synchronous (i.e., a class taught by an educator in real time) components¹. Nonadaptive e-learning environments, the most widespread type of e-learning environment today, provide a standardized training for all learners⁵⁶. While they can include instructional design variations (e.g., interactivity, feedback, practice exercises), they do not consider learners' characteristics and the data generated during the learning process to provide a personalized training⁶⁻⁸. This is problematic, since the interaction of health professionals and students with e-learning environments during the learning process generates a significant amount of data⁹. However, designers of e-learning environments and educators rarely make use of this data to optimize learning efficacy and efficiency⁹.

In recent years, educational researchers have strived to develop e-learning environments that take a data-driven and personalized approach to education¹⁰⁻¹³. E-learning environments that take into account each learner's interactions and performance level could anticipate what types of content and resources meet the learner's needs, potentially increasing learning efficacy and efficiency¹³. Adaptive e-learning environments (AEEs) were developed for this purpose. AEEs collect data to build each learner's profile (e.g., navigation behavior, preferences, knowledge), and use simple techniques (e.g., adaptive information filtering, adaptive hypermedia) to implement different

types of adaptivity targeting the content, navigation, presentation, multimedia, or strategies of the training to provide a personalized learning experience ¹¹¹². In the fields of computer science and educational technology, the term *adaptivity* refers to the process executed by a system based on ICTs of adapting educational curriculum content, structure or delivery to the profile of a learner¹⁴. Two main methods of adaptivity can be implemented within an AEE. The first method, *designed adaptivity*, is expert-based and refers to an educator who designs the optimal instructional sequence to guide learners to learning content mastery. The educator determines how the curriculum will adapt to learners based on a variety of factors, such as knowledge or response time to a question. This method of adaptivity is thus based on the expertise of the educator who specifies how technology will react in a particular situation on the basis of the "if THIS, then THAT" approach. The second method, *algorithmic adaptivity*, refers to use of algorithms to determine, for instance, the extent of the learner's knowledge and the optimal instructional sequence. Algorithmic adaptivity requires more advanced adaptivity techniques and learner-modelling techniques derived from the fields of computer science and artificial intelligence (e.g. Bayesian knowledge tracing, rule-based machine learning, natural language processing) 10 15-18.

127 The variability in the degree and the complexity of adaptivity within AEEs mirrors the adaptivity 128 that can be observed in non–e-learning educational interventions. Some interventions, like one-129 on-one human instruction and small-group classroom instruction, generally have a high degree of 130 adaptivity since the instructor can adapt his teaching to the individual profiles of learners and 131 consider their feedback ¹⁹. Other interventions, like large-group classroom instruction, generally 132 have a low degree of adaptivity to individual learners. In some interventions, like paper-based 133 instruction (e.g., handouts, textbooks), there is no adaptivity at all.

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AEEs have been developed and evaluated primarily in academic settings for students in mathematics, physics and related disciplines, for the acquisition of knowledge and development of cognitive skills (e.g., arithmetic calculation). Four meta-analyses reported on the efficacy of AEEs among high school and university students in in these fields of study ^{15-17 20}. The results are promising: AEEs are in almost all cases more effective than large-group classroom instruction. In addition, Nesbit, et al.²¹ point out that AEEs are more effective than nonadaptive e-learning environments. However, despite evidence of the efficacy of AEEs for knowledge acquisition and skill development in areas such as mathematics in high school and university students, their efficacy in improving learning outcomes in health professionals and students has not yet been established. To address this need, we conducted a systematic review and meta-analysis to identify and quantitatively synthesize all comparative studies of AEEs involving health professionals and students.

146 Systematic Review and Meta-Analysis Objective

147 To systematically identify, appraise, and synthesize the best available evidence regarding the
148 efficacy of AEEs in improving knowledge, skills, and clinical behavior in health professionals
149 and students.

150 Systematic Review and Meta-Analysis Questions

151 We sought to answer the following questions with the systematic review:

- 1. What are the characteristics of studies assessing an AEE designed for health professionals' and students' education?
- 2. What are the characteristics of AEEs designed for health professionals' or students' education?

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156 We sought to answer the following question with the meta-analysis:

3. What is the efficacy of AEEs in improving knowledge, skills, and clinical behavior in
 health professionals and students in comparison with nonadaptive e-learning
 environments, and non–e-learning educational interventions?

METHODS

We planned and conducted this systematic review following the Effective Practice and
Organization of Care (EPOC) Cochrane Group guidelines²², and reported it according to the
Preferred Reporting Items for Systematic review and Meta-Analysis (PRISMA) standards²³ (see
Supplementary File 1). We prospectively registered (International Prospective Register of
Systematic Reviews #CRD42017065585) and published the protocol of this systematic review²⁴.
Thus, in this paper, we present an abridged version of the methods with an emphasis on changes
made to the methods since the publication of the protocol.

Study Eligibility

We included primary research articles that assessed an AEE with licensed health professionals, students, trainees, and residents in any discipline. We defined an AEE as a computer-based learning environment which collects data to build each learner's profile (e.g., navigation behavior, individual objectives, knowledge), interprets these data through algorithms, and adapts in real-time the content (e.g., showing/hiding information), navigation (e.g., specific links and paths), presentation (e.g., page layout), multimedia presentation (e.g., images, videos), or tools (e.g., different set of strategies for different types of learners) to provide a dynamic and evolutionary learning path for each learner^{10 14}. We used the definitions of each type of adaptivity proposed by Knutov and colleagues¹². We considered for inclusion studies in which AEEs had

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designed or algorithmic adaptivity, and studies including a co-intervention in addition to adaptive e-learning (e.g. paper-based instruction). We considered for inclusion primary research articles in which the comparator was: 1) a nonadaptive e-learning environment; 2) a non-e-learning educational intervention: 3) another AEE with design variations. While included in the qualitative synthesis of the evidence for descriptive purposes, the third comparator was excluded from the meta-analysis. Outcomes of interest were knowledge, skills, and behavior^{25 26}, and were defined as follows: 1) knowledge: subjective (e.g., learner self-report) or objective (e.g., multiple-choice question knowledge test) assessments of factual or conceptual understanding; 2) skills: subjective (eg, learner self-report) or objective (eg, faculty ratings) assessments of procedural skills (e.g. taking a blood sample, performing CPR) or cognitive skills (e.g. problem-solving, interpreting radiographs) in learners; 3) behavior: subjective (eg, learner self-report) or objective (eg, chart audit) assessments of behaviors in clinical practice (such as test ordering)⁶. In terms of study design, we considered for inclusion all controlled, quantitative studies in accordaSsnce with the EPOC Cochrane Review Group guidelines²⁷. We excluded studies that: 1) were not published in English or French; 2) were non-experimental; 3) were not controlled; 4) did not report on at least one of the outcomes of interest in this review; 5) did not have a topic related to the clinical aspects of health. **Study Identification**

We previously published our search strategy²⁴. Briefly, we designed a strategy in consultation
with a librarian to search CINAHL, EMBASE, ERIC, PsycINFO, PubMed and Web of Science
for primary research articles published since the inception of each database up to February 2019.
The search strategy revolved around 3 key concepts: "adaptive e-learning environments", "health

200 professionals/students", and "effects on knowledge/competence (skills)/behavior" (see

201 Supplementary file 2). To identify additional articles, we hand-searched 6 key journals (e.g.,

202 British Journal of Educational Technology, Computers and Education) and the reference lists of

203 included primary research articles.

204 Study Selection

We worked independently and in duplicate (G.F. and M.-A.M.-C./T.M.) to screen all titles and abstracts for inclusion using the EndNote software V8.0 (Clarivate Analytics). We resolved disagreements by consensus. We then performed the full-text assessment of potentially eligible articles using the same methodology. Studies were included in the meta-analysis if they had a non-AEE control group and had no missing data.

210 Data Extraction

One review author (G.F.) extracted data from included primary research articles using a modified version of the data collection form developed by the EPOC Cochrane Review Group ²⁸. The main changes made to the extraction form were the addition of specific items relating to the AEE assessed in each study. Two review authors (T.M., M.-F.D.) validated the data extraction forms by reviewed the contents of each form against the data in the original article, adding comments when changes were needed. For all studies, we extracted the following data items if possible:

• *the population and setting*: study setting, study population, inclusion criteria, exclusion criteria;

• *the methods*: study aim, study design, unit of allocation, study start date and end date, and duration of participation;

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3 4	221	• <i>the participants</i> : study sample, withdrawals and exclusions, age, sex, level of instruction,
5 6	222	number of years of experience as a health professional, practice setting, and previous
7 8 9	223	experience using e-learning;
) 10 11	224	• <i>the interventions</i> : name of intervention, theoretical framework, statistical model/algorithm
12 13	225	used to generate the learning path, clinical topic, number of training sessions, duration of
14 15 16	226	each training session, total duration of the training, adaptivity subdomains (method, goals,
10 17 18	227	timing, factors, types), mode of delivery, presence of other educational interventions and
19 20	228	strategies;
21 22	229	• the outcomes: name, time-points measured, definition, person measuring, unit of
23 24 25	230	measurement, scales, validation of measurement tool;
26 27	231	• <i>the results</i> : results according to our primary (knowledge) and secondary (skills, behavior)
28 29	232	outcomes, comparison, time-point, baseline data, statistical methods used, and key
30 31 32	233	conclusions.
33 34 35	234	We contacted the corresponding authors of included primary research articles to provide us with
36 37	235	missing data.
38 39 40 41	236	Assessment of the Risk of Bias
42 43	237	We worked independently and in duplicate (G.F. and T.M./MF.D.) to assess the risk of bias of
44 45 46	238	included primary research articles using the EPOC risk of bias criteria, based upon the data
40 47 48	239	extracted with the data collection form ²⁸ . A study was deemed at high risk of bias if the
49 50	240	individual criterion "random sequence generation" was scored at "high" or at "unclear" risk of
51 52	241	bias.
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242 Data Synthesis

 First, we synthesized data qualitatively using tables to provide an overview of the includedstudies, and of the AEEs reported in these studies.

Second, using the Review Manager (RevMan) software V5.1, we first conducted a meta-analysis to quantitatively synthesize the efficacy of AEEs versus other educational interventions in improving all learning outcomes. We included studies in the meta-analysis if the comparator wasn't another AEE, if they were randomized, and if they reported outcome data. We then conducted meta-analyses with the same comparison for each outcome for which data from at least 2 studies were available (i.e., knowledge, skills). For randomized controlled trials (RCTs), we converted each post-test mean and standard deviation (SD) to a standardized mean difference ([SMD], also known as Hedges g effect size [ES]). For crossover RCTs, we used means pooled across each intervention. We pooled ESs using a random effects model. Statistical significance was defined by a two-sided alpha of .05.

We first assessed heterogeneity qualitatively by examining the characteristics of included studies, the similarities and disparities between the types of participants, the types of interventions, and the types of outcomes. We then used the l^2 statistic within the RevMan software to quantify how much the results varied across individual studies (i.e., between-study inconsistency, or heterogeneity). We interpreted the I² values as follows: 0%-40%: might not be important; 30%-60%: may represent moderate heterogeneity; 50%-90%: may represent substantial heterogeneity; and 75%-100%: considerable heterogeneity²⁹. We performed sensitivity analysis to assess if the exclusion of studies at high risk of bias or with a small sample size (n < 20) would have had an

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63 impact on statistical heterogeneity. Subgroup analyses were performed to examine if study 64 population and study comparators were potential effect modifiers. 65 Since less then 10 studies were included in the meta-analysis for each outcome, we did not assess reporting biases using a funnel plot, as suggested in the Cochrane Handbook³⁰. 66 67 Assessment of the Quality of Evidence We worked independently and in duplicate (G.F. and M.-A.M.-C.) to assess the quality of 68 69 evidence for each individual outcome. We used the Grading of Recommendations Assessment, 70 Development, and Evaluation (GRADE) Web-based software, based upon the data extracted with the data collection checklist ³¹. We considered 5 factors (risk of bias of included studies, 71 72 indirectness of evidence, unexplained heterogeneity or inconsistency of results, imprecision of 73 the results, probability of reporting bias) for downgrading the quality of the body of evidence for ie. each outcome 31 . 74

75 **Patient and Public Involvement**

76 Patients and the public were not involved in setting the research question, the outcome measures, 77 the design or conduct of this systematic review. Patients and the public were not asked to advise 78 on interpretation of results or to contribute to the writing or editing of this document.

- RESULTS 79
 - 80 **Study Flow**

81 From a pool of 10,569 potentially relevant articles, we found 21 quantitative, controlled studies 82 assessing an AEE with health professionals or students (see Figure 1).

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[Insert Figure 1]

Out of 21 included studies in the qualitative synthesis, 4 studies compared two AEEs with design variations³²⁻³⁵, and 4 studies had missing data³⁶⁻³⁹. The 4 studies with missing data did not report properly data regarding the results, i.e. mean scores and standard deviations in both study groups at post-test, regarding the outcomes of interest in this review (i.e., knowledge, skills or clinical behavior). Thus, 13 studies were included in the meta-analysis and used to calculate an ES on learning outcomes.

290 Study Characteristics

We summarized the key characteristics of included studies in table format (see **Table 1**). In terms of study population, in the 21 studies examined published between 2003 and 2018, investigators have evaluated AEEs mostly in the medical field. Studies focused on medical students (n=8) ³⁸⁻⁴⁴, medical residents (n=8) 32-354145-47, physicians in practice (n=4) 36374148, nursing students (n=2) 4049 , nurses in practice (n=2) 4850 and health sciences students (n=1) 51 . Three studies focused on multiple populations ^{40 41 48}. The median sample size was 46 participants (interguartile range [IQR] 123). In terms of study design, 15 out of 21 studies (71%) were randomized, 7 studies of which were randomized crossover trials ^{33 34 41-43 45 47}. The median number of training sessions was 2 (IQR 2.5 sessions), the median training time was 2.13 hours (IQR 2.88 hours), and the median training period was 14 days (IQR 45 days). In terms of comparators, it is possible to underline three types of comparisons. The first comparison is an AEE versus another AEE with design variations $(n=4)^{32-35}$, which implies that one of the AEEs assessed had variations in its design, such as different types of adaptivity (e.g., feedback in one AEE is longer or more complex than in the other). The second comparison is an AEE versus a nonadaptive e-learning

305	environment (n=11) ^{38-40 42-46 48 50 51} . The third and final comparison is an AEE versus another
306	type of educational intervention, such as a paper-based educational intervention, including
307	handouts, textbooks or images $(n=3)^{37 41 47}$, or a traditional educational intervention, such as a
308	group lecture (n=2) $^{49.52}$. In one study, the comparator was not clearly reported 36 . As stated
309	before, only the second and third types of comparisons were included in the meta-analysis since
310	our aim was to synthesize quantitatively the efficacy of AEEs versus other types of educational
311	interventions.

Finally, regarding the outcomes, knowledge was assessed in 14 out of 21 studies (66.7%) ^{32 35 36 38} $^{39\,41-45\,47-50}$, skills in 9 studies (42.9%) $^{33\,34\,36\,37\,40\,46\,50-52}$, and clinical behavior in 2 studies (9.5%) ^{44 50}. Outcome measures for knowledge were similar across studies: in 9 out of 14 studies measuring knowledge, investigators employed multiple-choice questionnaires developed by the research team with input from content experts that were tailored to training content to ensure specificity. Knowledge was also assessed using true-false questions in two studies, and the type of questionnaire was not specified in three studies. Outcome measures for skills were also similar across the 9 studies reporting this outcome, since in all studies investigators measured *cognitive* skills rather than *procedural* skills. Indeed, all outcomes measures for skills were related to clinical reasoning. In 6 studies, skills were measured through tests that included a series of diagnostic tests (eg electrocardiograms, x-rays, miscroscopy images) that learners had to interpret. In 3 studies, skills wre measured through questions based on clinical situations in which learners had to specify how they would react in these particular situations. We were not able to describe the similarity between the outcome measures for clinical behaviour no details were provided in one of the two studies reporting this outcome.

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Table 1. Characteristics of Included Studies.

First Author, Year Country	Participants*	Study Design [†]	No. and Duration of Training Sessions	Duration of Intervention	Comparison(s) [‡]	Type of Outcome(s) of Interest [§]	Outcome Measures
Comparison: a	daptive e-learni	ng environments vs. other educa	ational interventions				
Casebeer, 2003	PP; N = 181	RCT; posttest-only, 2 groups	4 sessions; 1 hour each	NR	NR	Knowledge	21-item multiple-choice
USA Cook, 2008 USA	R; N = 122	RXT; posttest-only, 4 groups	4 sessions; 30 minutes each	126 days	NEE	Skills Knowledge	69-item case-based multiple- choice questionnaire
Crowley, 2010 USA	PP; N = 15	RCT; pretest–posttest, 2 groups	4 sessions; 4 hours each	138 days	Р	Skills	Virtual slide test to examine diagnostic accurary
de Ruijter,			Nie Fund energinge	100 dava		Knowledge	18-item true-false questionnaire (range 0–18)
2018 Netherlands	NP; N = 269	RCI; pretest-posttest, 2 groups	No fixed sessions	180 days	NEE	Behavior	9-item self-reported questionna (range 0–9)
Hayes-Roth, 2010 USA	MS, NS; N = 30	RCT; pretest–posttest– retention-test, 3 groups	NR; mean training time 2.36 hours	NR	1. NEE 2. NI	Skills	6-item written skill probe (range 6–18)
						Knowledge	Unclear
Lee, 2017 USA	MS; N = 1522	NRCT; pretest-posttest, 3 groups	5 sessions; NR	42 days	NEE	Skills	Multidimensional situation-base questions –RealIndex (range 0–100%)
						Behavior	Unclear
Micheel, 2017 USA	PP, NP; N = 751	NRCT; pretest–posttest– retention-test, 2 groups	NR	NR	NEE	Knowledge	10-item true-false questionnaire (range 0–10)
Morente, 2013 Spain	NS; N = 73	RCT; pretest-posttest, 2 groups	1 session; 4 hours	1 day	Т	Knowledge	22-item multiple-choice questionnaire (range 0–22)
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Munoz, 2010 Colombia	MS; N = 40	NRCT; pretest–posttest, 2 groups	NR; mean training time 5.97 hours	NR	NEE	Knowledge	10-item multiple-choice questionnaire (range 0–10)
Romito, 2016 USA	R; N = 24	NRCT; pretest–posttest– retention-test, 2 groups	1 session; 30 minutes	1 day	NEE & T	Skills	22-item videoclip-based test
Samulski, 2017 USA	MS, R, PP; N = 36	RXT; pretest–posttest, 2 groups	2 sessions; 20 minutes to 14 hours	1 month	Ρ	Knowledge	28-item multiple-choice questionnaire (range 0–100%)
Thai, 2015 USA	HSC; N = 87	RCT; pretest–posttest– retention-test, 3 groups	1 session; 45 minutes	1 day	1. AEE 2. NEE	Skills	14-item case-based test (range 0- 100%)
Van Es, 2015 Australia	R; N = 43	RXT; posttest-only, 2 groups	3 sessions; NR	50 days	Р	Knowledge	7 to 21-item multiple-choice questionnaire (range 0–100%)
Van Es, 2016 Australia	MS; N = 46	RXT; posttest-only, 2 groups	3 sessions; 2 hours each	34 days	NEE	Knowledge	Multiple-choice questionnaire
Wong, 2015 Australia	MS; N = 99	RXT; posttest-only, 2 groups	2 sessions; 1.5 hour each	14 days	NEE	Knowledge	8-item multiple-choice and interactive questions (range 0–100%)
Wong, 2017 USA	MS; N = 178	NRCT; pretest–posttest– retention-test, 3 groups	1 session; NR	35 days	1. T 2. AEE & T	Skills	Test to examine diagnostic accuracy
Woo, 2006 USA	MS; N = 73	NRCT; pretest–posttest, 3 groups	1 session; 2 hours	1 day	1. NEE 2. NI	Knowledge	Short-response questionnaire
Comparison: a	idaptive e-learn	ing vs. adaptive e-learning (two A	AEEs with design variations)				
Crowley, 2007 USA	R; N = 21	RCT; pretest–posttest– retention-test, 2 groups	1 session; 4.5 hours	1 day	AEE	Knowledge	51-item multiple-choice questionnaire (range 0–100%)
El Saadawi, 2008 USA	R; N = 20	RXT; pretest–posttest, 2 groups	2 sessions; 2 hours each	1 day	AEE	Skills	Virtual slide test to examine diagnostic accuracy
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El Saadawi, 2010 USA	R; N = 23	RXT; pretest–posttest, 2 gro	ups 2 sessions; 2.25 hours each	2 days	AEE	Skills	Virtual slide test to examine diagnostic accuracy
Feyzi- Begnagh, 2014 USA	R; N = 31	RCT; pretest–posttest, 2 gro	ups 2 sessions; 2 & 3 hours	1 day	AEE	Knowledge	Unspecified test
* Participants: students. † Study desigr ‡ Comparison	: MS indicates me n: RCT indicates : AEE indicates a	edical students; NS, nursing stud randomized controlled trial; RXT idaptive e-learning environment;	ents; R, residents (physicians in po randomized crossover trial; NRCT NEE, nonadaptive e-learning envir	ostgraduate tra 「, non-random onment; NI, no	ining); PP, physiciar zed controlled trial. intervention contro	ns in practice; NP, nurse I group; T, traditional (g	es in practice; HSC, health sciences roup lecture); P, paper (handout, textboo
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334 Characteristics of Adaptive E-Learning Environments

335 We summarized the key characteristics of AEEs assessed in the 21 studies in table format (see 336
 Table 2). In terms of the clinical topics of the AEEs, the majority of AEEs focused on training
 337 medical students and residents in executing and/or interpreting diagnostic tests. Indeed, a 338 significant proportion of the AEEs assessed focused on dermopathology and cytopathology 339 microscopy ^{32-35 37 41 42 47} (n=8). Other topics were diagnostic imaging ^{43 46} (n=2), behavior change counseling ^{40 50} (n=2), chronic disease management ^{45 48} (n=2), pressure ulcer evaluation ⁴⁹ (n=1), 340 childhood illness management 38 (n=1), electrocardiography 51 (n=1), fetal heart rate 341 342 interpretation ⁵² (n=1), hemodynamics ³⁹ (n=1), chlamydia screening (n=1) ³⁶ and atrial fibrillation management $(n=1)^{44}$. 343 344 The 21 AEEs examined were based on a wide variety of theoretical frameworks. The most 345 frequently used framework was cognitive tutoring, adopted in 5 studies ^{32-35 37}, which refers to the

346 use of a cognitive model. The integration of a cognitive model in an AEE implies the 347 representation of all the knowledge in the field of interest in a way that is similar to the human 348 mind for the purpose of understanding and predicting the cognitive processes of learners ⁵³. The second most used framework was perceptual learning, adopted in 3 studies ^{46 51 52}. Perceptual 349 350 learning aims at improving information extraction skills of the environment and the development 351 of automaticity in this respect in learners ⁴⁶. Interestingly, 2 studies used models from behavioral science, the Transtheoretical Model ³⁶ and the I-Change Model ⁵⁰, to tailor the AEE to the 352 353 theoretical determinants of clinical behavior change in nurses and physicians in practice. 354 Theoretical frameworks relating to self-regulated learning ³⁵, learning styles ^{38 48}, guided mastery ⁴⁰, and cognitive load ⁴³, problem-based-learning ³⁶, and situated learning ³⁶ were also used. 355

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356	Three main adaptive e-learning platforms were used by investigators in studies examined:
357	SlideTutor (n=4) ^{33 37 54 55} , Smart Sparrow (n=4) ^{41-43 47} , and the Perceptual Adaptive Learning
358	Module (PALM, n=3) ^{51 52 56} . SlideTutor is an AEE with algorithmic adaptivity which provides
359	cases to be solved by learners under supervision by the system. These cases incorporate
360	dermopathology virtual slides that must be examined by learners to formulate a diagnosis. An
361	expert knowledge base, consisting of evidence-diagnosis relationships, is used by SlideTutor to
362	create a dynamic solution graph representing the current state of the learning process and to
363	determine the optimal instructional sequence ⁵⁵ . Smart Sparrow is an AEE with designed
364	adaptivity which allows educators to determine adaptive factors, such as answers to questions,
365	response time to a question, and learner actions, to specify how the system will adapt the
366	instructional sequence or provide feedback. These custom learning paths can be more or less
367	personalized ⁴² . PALM is an AEE with algorithmic adaptivity aiming to improve perceptual
368	learning through adaptive response-time-based sequencing to determine dynamically the spacing
369	between different learning items based on each learner's accuracy and speed in interactive
370	learning trials ⁵¹ . Different custom adaptive e-learning platforms were used in other studies.
371	

Table 2. Characteristics of adaptive e-learning environments.

First Author, Year		Theoretical Framework(s)	Platform	Adaptivity Subdomains					
	Clinical Topic(s)			Adaptivity Method	Adaptivity Goals	Adaptivity Timing	Adaptivity Factors	Adaptivity Ty	
Casebeer, 2003	Chlamydia screening	Transtheoretical model of change; Problem-based learning; Situated learning theory	NR	Designed Adaptivity	To increase learning effectiveness (knowledge, skills).	Throughout the training, after case- based and practice- based questions.	User answers to questions	ContentNavigation	
Cook, 2008	Diabetes, hyperlipidemia, asthma, depression	NR	NR	Designed Adaptivity	To increase learning efficiency (knowledge gain divided by learning time).	After each case- based question in each module (17 to 21 times/module).	User knowledge	ContentNavigation	
Crowley, 2007	Dermopathology; subepidermal vesicular dermatitis	Cognitive Tutoring	SlideTutor	Algorithmic Adaptivity	To increase learning gains, metacognitive gains, and diagnostic performance.	At the beginning of each case.	User actions: results of problem- solving tasks; requests for help	 Content Navigation Presenta Multimed Tools 	
Crowley, 2010	Dermopathology; melanoma	Cognitive Tutoring	SlideTutor	Algorithmic Adaptivity	To improve reporting performance and diagnostic accuracy.	At the beginning of each case.	User actions: results of problem- solving tasks; reporting tasks; requests for help	 Content Navigation Presenta Multimed Tools 	
de Ruijter, 2018	Smoking cessation counseling	I-Change Model	Computer- Tailored E- Learning Program	Designed Adaptivity	To modify behavioral predictors and behavior.	At the beginning of the training.	Demographics, behavioral predictors, behavior	Content	
El Saadawi, 2008	Dermopathology; melanoma	Cognitive Tutoring	ReportTutor	Algorithmic Adaptivity	To teach how to correctly identify and document all relevant	At the beginning of each case.	User actions, report features	 Content Navigation Presenta Multimed 	

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Micheel, 2017	Oncology	Learning Style Frameworks	Learning- style tailored educational platform	Designed Adaptivity	To increase learning effectiveness (knowledge).	After assessing the learning style.	Learning style	• • •	⊃resentation Multimedia Tools
Lee, 2017	Treatment of atrial fibrillation	NR	Learning Assessment Platform	Designed Adaptivity	To increase learning effectiveness (knowledge, competence, confidence and practice).	After learning gaps identified in the first session.	Learning gaps in relation to objectives	• (Content
Hayes- Roth, 2010	Brief intervention training in alcohol abuse	Guided Mastery	STAR Workshop	NR	To improve attitudes and skills.	During clinical cases.	User scores, user- generated dialogue	• (Content Navigation
Feyzi- Begnagh, 2014	Dermopathology; nodular and diffuse dermatitis	Cognitive Tutoring, Theories of Self-Regulated Learning	SlideTutor	Algorithmic Adaptivity	To improve metacognitive and learning gains during problem solving.	During each case or immediately after each case.	User actions: results of problem- solving tasks; reporting tasks; requests for help	• (• • •	Content Navigation Presentation Multimedia Tools
El Saadawi, 2010	Dermopathology	Cognitive Tutoring	SlideTutor	Algorithmic Adaptivity	To facilitate transfer of performance gains to real- world tasks that do not provide direct feedback on intermediate steps.	During intermediate problem-solving steps.	User actions: results of problem- solving tasks; reporting tasks; requests for help	• (• •	Content Navigation Presentatio Multimedia
					in the diagnostic report.				

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Morente, 2013	Pressure ulcer evaluation	NR	ePULab	Designed Adaptivity	learning effectiveness (knowledge, skills).	Each pressure ulcer evaluation.	User skills	• Cor
Munoz, 2010	Management of childhood illness	Learning Styles Framework	SIAS-ITS	Designed Adaptivity	To increase learning effectiveness and efficiency.	At the beginning of the training.	User knowledge, user learning style	CorToc
Romito, 2016	Transoesophageal echocardiography	Perceptual Learning	TOE PALM	Algorithmic Adaptivity	To improve response accuracy and response time.	After each clinical case.	User response accuracy, user response time	CorNavMul
Samulski, 2017	Cytopathology; pap test, squamous lesions, glandular lesions	NR	Smart Sparrow	Designed Adaptivity	To improve learning effectiveness.	During intermediate problem-solving steps.	User knowledge	CorNav
Thai, 2015	Electrocardiography	Perceptual Learing Theory; Adaptive response-time based algorithm	PALM	Algorithmic Adaptivity	To improve perceptual classification learning effectiveness and effiency.	After each user response.	User response accuracy, user response time	 Cor Pre Mul Too
Van Es, 2015	Diagnostic cytopathology; gynecology, fine needle aspiration, exfoliative fluid	NR	Smart Sparrow	Designed Adaptivity	To improve learning effectiveness.	During intermediate problem-solving steps.	User responses	 Cor Nav Pre Mul
Van Es, 2016	Diagnostic cytopathology; gynecology, fine needle aspiration, oxfoliativo fluid	NR	Smart Sparrow	Designed Adaptivity	To improve learning effectiveness.	During intermediate problem-solving steps.	User responses	Cor Nav Pre Mul

	Wong, 2015	Diagnostic imaging; chest X-rays, CT scans	Cognitive Load Theory	Smart Sparrow	Designed Adaptivity	To improve learning effectiveness.	During intermediate problem-solving steps.	User responses	٠	Content
	Wong, 2017	Fetal heart rate interpretation	Perceptual Learning	PALM	Algorithmic Adaptivity	To improve response accuracy and response time.	After each clinical case.	User response accuracy, user response time	• •	Content Navigation Multimedia
	Woo, 2006	Hemodynamics; baroreceptor reflex	NR	CIRCSIM- Tutor	Algorithmic Adaptivity	To improve knowledge related to problem- solving tasks.	After each user response.	User knowledge, user responses	• •	Content Navigation Tools
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We propose 5 subdomains that emerged from the review to characterize the adaptivity process of
AEEs reported in the 21 studies: adaptivity method, adaptivity goals, adaptivity timing,
adaptivity factors and adaptivity types.

380 First Subdomain: Adaptivity Method

This subdomain relates to the method of adaptivity that dictates how the AEE adapts instruction to a learner. As we previously described, there are two main methods of adaptivity: *designed adaptivity* and *algorithmic adaptivity*. The first is based on the expertise of the educator who specifies how technology will react in a particular situation on the basis of the "if THIS, then THAT" approach. The second refers to use of algorithms that will determine, for instance, the extent of the learner's knowledge and the optimal instructional sequence. In this review, 11 AEEs employed designed adaptivity ^{36 38 41-44 47-50 57}, and 9 AEEs employes algorithmic adaptivity ^{33 37 39} ^{51 52 54-56 58}. The adaptivity method wasn't specified in one study ⁴⁰.

3 389 Second Subdomain: Adaptivity Goals

This subdomain relates to the purpose of the adaptivity process within the AEE. For most AEEs, the adaptivity process aims primarily to increase the efficacy and/or efficiency of knowledge acquisition and skills development relative to other training methods ^{32 35 36 38-45 47-49 51}. For instance, several AEEs aimed to increase the diagnostic accuracy and reporting performance of medical students and residents ^{32-34 37 46 52}. In one study, the goal of adaptivity was to modify behavioral predictors and behavior in nurses ⁵⁰. In cases where two adaptive AEEs with certain variations in their techno-pedagogical design are compared with each other, the adaptivity process generally aims at improving the metacognitive and cognitive processes related to learning ^{32 33 35}.

Third Subdomain: Adaptivity Timing

400 This subdomain relates to *when* the adaptivity occurs during the learning process with the AEE.

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401 In 19 out of 21 studies, the adaptivity occurred throughout the training with AEE, usually after an

402 answer to a question or during intermediate problem-solving steps. However, in two studies,

403 adaptivity was only implemented at the beginning of the training with the AEE following survey

404 responses 3850 .

405 Fourth Subdomain: Adaptivity Factors

406 This subdomain relates to the learner-related data (variables) upon which the adaptivity process is
407 based. The most frequently targeted variable is the learner's scores after an assessment or a
408 question within the AEE (e.g., knowledge/skills scores, response accuracy scores) ³⁸⁻⁴³ ⁴⁵⁻⁴⁷ ⁴⁹ ⁵¹ ⁵².
409 Other frequently targeted variables include the learner's actions during its use of the AEE (e.g.,
410 results of problem-solving tasks, results of reporting tasks, requests for help) ³²⁻³⁵ ³⁷, and the

411 learner's response time regarding a specific question or task ^{46 51 52}.

412 Fifth Subdomain: Adaptivity Types

The final subdomain relates to which types of adaptivity are mobilized in the AEE: content, navigation, multimedia, presentation and tools. In the context of this review, the adaptivity types are based upon the work of Knutov and colleagues ¹². Overall, 17 out of 21 (81%) AEEs examined integrated more than one type of adaptivity. Content adaptivity was the most used adaptivity type; it was implemented in all but one AEEs reviewed (n=20). Content adaptivity aims to adapt the textual information (curriculum content) to the learner's profile through different mechanisms and to different degrees ¹². Navigation adaptivity was the second most used adaptivity type (n=14). Navigation can be adapted in two ways; it can be enforced or suggested.

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When enforced, an optimal personalized learning path is determined for the learner by an expert educator or by the algorithms within the AEE. When suggested, there are several personalized learning paths available to each learner, who can determine the path he prefers himself¹². Most reviewed studies included AEEs with enforced navigation, with one optimal personalized learning path being determined by an expert educator or by the algorithm. *Multimedia adaptivity* was the third most used adaptivity type (n=11). This adaptivity type, much like content adaptivity which relates to textual information, implies the adaptivity of the multimedia elements of the training such as videos, pictures, models, to the learner's profile. *Presentation adaptivity* was the fourth most used adaptivity type (n=9). It implies the adaptivity of the layout of the page to the digital device used, or to the learner's profile. *Tools adaptivity* was the least used adaptivity type (n=8). This technique results in providing a different set of features or learning strategies for different types of learners, such as different interfaces for problem solving, and knowledge 1.04 representation.

Risk of Bias Assessment

Results of included studies for the risk of bias assessment are presented in Figure 2 and Figure 3. In \geq 75% of studies, biases related to similarity of baseline outcome measurements, blinding of outcome assessment and selective reporting of outcomes were low. Moreover, in \geq 50% of studies, biases related to contamination were low. Regarding the blinding of outcome assessment, in most studies, review authors judged that the outcomes of interest and the outcome measurement were not likely to be influenced by the lack of blinding, since studies had objective measures, i.e. an evaluative test of knowledge or skills. Regarding contamination bias, review authors scored studies at high risk if they had a crossover design.

1 2								
2 3 4	443	However, in \geq 50% of studies, biases related to random sequence generation, allocation						
5 6 7 8 9 10 11	444	concealment, similarity of baseline characteristics, similarity of baseline characteristics, blinding	,					
	445	of participants and personnel, and incomplete outcome data were unclear or high. Regarding						
	446	random sequence generation, an important number of studies did not report on the method of						
12 13	447	randomization used by investigators. As per Cochrane recommendations, all eligibile studies						
14 15	448	were incuded in the meta-analysis, regardless of the risk of bias assessment. Indeed, since almost	t					
16 17 18	449	all studies scored overall at unclear risk of bias, Cochrane suggests to present an estimated						
19 20	450	intervention effect based on all available studies, together with a description of the risk of bias in	l					
21 22	451	individual domains ³⁰ .						
23 24								
24 25 26 27 28	452 [Insert Figure 2]							
	453	[Insert Figure 3]						
29 30								
30 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47	454	Quantitative Results						
	455	Efficacy of AEEs versus other educational interventions in improving knowledge						
	456	The pooled ES (standardized mean difference [SMD] 0.70; 95% confidence interval [CI] -0.08-						
	457	1.49; Z =1.76, P 0.08) of AEEs compared to other educational interventions in improving						
	458	knowledge suggests a medium to large effect (see Figure 4). However, this result is not						
	459	statistically significant. Significant statistical heterogeneity was observed among studies (I^2						
	460	=97%, P <.00001), and individual ESs ranged from -1.10 to 3.05. One study in particular 45						
48 49	461	reported a negative ES, but the difference between groups in knowledge scores was statistically						
50 51 52	462	nonsignificant. Moreover, while participants using the AEE in the experimental group reported						
53 54	463	the same knowledge scores as participants in the control group at the end of study, time spent on						
55 56	464	instruction was reduced by 18% with the AEE compared to the nonadaptive e-learning						
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59 60		For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml	3					

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2 3 4	465	environment, thus improving learning efficiency ⁴⁵ . When that study ⁴⁵ is removed from the meta-						
5 6	466	analysis, the pooled ES becomes statistically significant (SMD 1.07; 95% CI 0.28-1.85; $Z = 2.67$,						
7 8	467	<i>P</i> 0.008).						
9 10 11 12	468	[Insert Figure 4]						
13 14 15 16 17	469	Efficacy of AEEs versus other educational interventions in improving skills						
	470	As we considered ESs larger than 0.8 to be large ⁵⁹ , the pooled ES (SMD 1.19; 95% CI 0.59-						
19 20	471	1.79; Z =3.88, P 0.0001) of AEEs compared to other educational interventions in improving						
21 22	472	skills suggests a significantly large effect (see Figure 5). Statistical heterogeneity was lower than						
23 24	473	in previous analyses, but was still significant ($I^2 = 89\%$, P <.00001). Individual ESs ranged from						
25 26 27 28 29 30 31 32 33 34 35	474	0.17 to 2.87.						
	475	[Insert Figure 5]						
	476	For both knowledge and skills, we conducted subgroup analyses according to population (health						
	477	professionals versus students) and comparator (adaptive e-learning versus nonadaptive e-						
30 37 38	478	learning, adaptive e-learning versus paper-based instruction, adaptive e-learning versus						
39 40	479	classroom-based instruction). No statistically significant differences between subgroups were						
41 42 43	480	found regarding the effect sizes.						
44 45 46	481	Quality of the Evidence						
47 48 49	482	The quality of evidence table produced with GRADE, as well as the justifications for each						
50 51	483	decision, is presented in Supplementary File 3 (GRADE quality of evidence levels: very low, low,						
52 53	484	moderate, high). For knowledge, the quality of evidence was deemed to be very low. More						
54 55 56	485	precisely, risk of bias was deemed serious, inconsistency serious, indirectness not serious, and						
57 58		3						
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486 imprecision serious. For skills, the quality of evidence was deemed to be low. More precisely,
487 risk of bias was deemed serious, inconsistency serious, indirectness not serious, and imprecision
488 serious.

DISCUSSION

490 Principal Findings

This is the first systematic review and meta-analysis to evaluate the efficacy of AEEs in health professionals and students. We identified 21 relevant studies published since 2003, 17 of which assessed an AEE versus another educational intervention (large-group classroom instruction, nonadaptive e-learning environment or paper-based learning), and 4 of which assessed 2 AEEs with design variations head-to-head. When compared with other educational interventions, AEEs were associated with statistically significant improvements in learning outcomes in 12 out of 17 studies. Pooled ESs were medium to large for knowledge and large for skills, but only the latter was associated with a statistically significant effect. Statistical heterogeneity was high in all analyses. However, this finding is consistent with other meta-analyses in the field of medical education that also reported high heterogeneity across studies ^{8 60 61}. No potential effect modifiers were found during subgroup analyses, and these did not help in explaining the source of the heterogeneity. The quality of evidence for all comparisons was either low or very low. Therefore, while we believe the results support the potential of AEEs for the education of health professionals and students, we recommend interpreting the ESs with caution.

505 Comparison with Other Studies

506 To our knowledge, no previous systematic review and meta-analysis has specifically assessed the 507 efficacy of AEEs in improving learning outcomes in health professionals and students, or any Page 33 of 54

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other population. However, interestingly, since the 1990's there has been a strong research interest in the field of AEEs with algorithmic adaptivity (also known as intelligent learning environments [IEEs] or intelligent tutoring systems [ITSs]) into elementary, high school and postsecondary education for multiple subjects ¹⁷. Thus, multiple meta-analyses have been conducted with regard to AEEs in that setting. Steenbergen-Hu and Cooper ¹⁵ reported a mean ES of 0.35 of AEEs with algorithmic adaptivity on learning outcomes in college students when compared to all other types of educational interventions. The mean ES was 0.37 when the comparator was large-group classroom instruction, 0.35 when the comparator was nonadaptive e-learning, and 0.47 when the comparator was textbooks or workbooks¹⁵. Ma, et al. ¹⁸ reported a mean ES of 0.42 of AEEs with algorithmic adaptivity on learning outcomes in elementary, high school and postsecondary students when compared to large-group classroom instruction. The mean ES was 0.57 when the comparator was nonadaptive e-learning, and 0.35 when the comparator was textbooks or workbooks. Interestingly, the mean ES was

higher for studies which assessed an AEE in biology and physiology (0.59) and in humanities and
social science (0.63) than in studies which assessed an AEE in mathematics (0.35) and physics
(0.38)¹⁸.

Kulik and Fletcher ¹⁷ reported a mean ES of 0.65 of AEEs with algorithmic adaptivity on
learning outcomes in elementary, high school, and postsecondary students when compared to
large-group classroom instruction. Education areas in this review were diverse (e.g.,
mathematics, computer science, physics), but none were related to health sciences. Interestingly,
the mean ES was 0.78 for studies up to 80 participants, and 0.30 for studies with more than 250

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participants. Moreover, the mean ES for studies conducted with elementary and high school
students was 0.44, compared to 0.75 for studies conducted with postsecondary students¹⁷.

532 Thus, in light of the results of these meta-analyses, the ES reported in our review may appear 533 high. However, our review looked more specifically into the efficacy of AEEs in improving 534 learning outcomes in health professionals and students. This is significant since, in the meta-535 analyses of Steenbergen-Hu and Cooper¹⁵, Ma, et al.¹⁸, and Kulik and Fletcher¹⁷, AEEs seem to be more effective in postsecondary students ^{17 18} and for learning subjects related to biology, 536 physiology and social science ¹⁸. Moreover, previous meta-analyses focused on the efficacy of 537 538 AEEs in improving procedural and declarative knowledge, and did not report on the efficacy on 539 AEEs in improving skills. This is important since AEEs may be more effective for providing 540 tailored guidance and coaching for developing skills regarding complex clinical interventions, rather than learning factual knowledge, which generates less cognitive load ^{62 63}. 541

542 Implications for Practice and Research

543 This review provides important implications for the design and development of AEEs for health 544 professionals and students. Table 3 presents 8 practical considerations for the design and 545 development of adaptive e-learning environments based on the results of this systematic review 546 for educators and educational researchers.

547Table 3.Practical considerations for the design and development of adaptive e-learning548environments.

Practical considerations	Explanations
Developing the Educational Content	• Given the adaptivity and the different learning pathways inherent to adaptive e- learning environments (AEEs), it is necessary to develop more pedagogical content (e.g. 60 minutes of learning) to reach the planned duration of each adaptive e-learning session (e.g. 30 minutes of learning).

	Selecting a Theoretical Framework	• Selecting a theoretical framework coherent with the underlining principles of adaptivity of AEEs is crucial. These frameworks can be related to human cognition (e.g. Cognitive Load Theory, Cognitive Tutoring), behavior change (e.g. Transtheoretical model, I-Change Model) or learning (e.g. Perceptual Learning, Situated Learning).	
	Selecting the Adaptivity Method	 Selecting the adaptivity method refers to how the AEE will adapt its instructional sequence. There are two main adaptivity methods: Designed adaptivity is based on the expertise of the educator who designs personalized pathways to guide learners to learning content mastery; Algorithmic adaptivity is based on different algorithms to determine, for instance, the extent of the learner's knowledge and the optimal instructional pathway. 	-
	Selecting the Adaptivity Goal(s)	• Selecting the adaptivity goal(s) is important, since it will dictate how the instruction will be adapted in the AEE. The goal of adaptivity within an AEE may be to increase learning effectiveness, increase learning efficiency, modify behavioral predictors, or improve cognitive/metacognitive processes related to learning.	
	Selecting the Adaptivity Timing	• Selecting the timing of adaptivity within an AEE relates to <i>when</i> the adaptivity occurs during the learning process. Adaptivity can be implemented at the beginning of the training only, or throughout the training. Adaptivity timing is closely linked to which adaptivity factor(s) are targeted in learners.	
	Selecting the Adaptivity Factor(s)	• Adaptivity factors are essentially data upon which the adaptivity process is based. These data can be related to the learner's performance (e.g. knowledge, skills), his behavior / actions on the page (e.g. response time, requests for help), his overall learning path on the platform, or any other variables of interest in the learner.	
	Selecting the Adaptivity Type(s)	 Multiple types of adaptivity can be implemented in an AEE: Content adaptivity refers to the adaptation of the textual information; Navigation adaptivity refers to the adaptation of the curriculum sequence; Presentation adaptivity refers to the adaptation of layout of the screen to the digital device used, or to the learner's profile; Multimedia adaptivity refers to the adaptation of multimedia elements of the training such as videos, pictures, models; Tools adaptivity refers to the adaptation of training features, learning strategies or learning assessment methods (e.g. interface for problem solving). 	
	Determining your technical resources and selecting the adaptive e-learning platform	• After the content has been developed, the theoretical framework has been selected and the decisions related to the different subdomains adaptivity have been made, it is crucial to determine your technical resources and evaluate preexisting adaptive elearning software to determine if it meets your needs and goals. If you plan to employ a specialist or team to develop the platform, estimate development cost and timeline.	
549	This review also pro	vides several key insights for future research. In terms of <i>population</i> , future	-
550	research should focu	is on assessing AEEs with health professionals in practice, such as registered	
551	nurses and physician	is, rather than students in these disciplines. This could provide key insights	
552	into how AEEs can	impact clinical behavior and, ultimately, patient outcomes. In addition,	
553	invetsigators should	target larger sample sizes. In terms of <i>interventions</i> , researchers should	
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554 report more clearly on adaptivity methods, goals, timing, factors and types. Moreover, 555 researchers should provide additional details regarding the underlining algorithms allowing the 556 adaptivity process in order to ensure replicability of findings. Regarding *comparators*, this review 557 suggests there is a need for additional research using traditional comparators (i.e., large group 558 classroom instruction) and more specific comparators (i.e., adaptive e-learning environment with 559 design variations). Regarding *outcomes* and outcome measures, researchers should use validated 560 measurement tools of knowledge, skills, and clinical behavior to facilitate knowledge synthesis. 561 Moreover, the very low number of studies assessing the impact of AEEs on health professionals' 562 and students' clinical behavior demonstrates the need for further research with higher-level 563 outcomes. Finally, in terms of *study designs*, researchers should focus on research designs 564 allowing the assessment of the impact of multiple educational design variations and adaptivity 565 types within one study, such as factorial experiments. 4

566 **Strengths and Limitations**

Strengths of this systematic review and meta-analysis include the prospective registration and 567 568 publication of a protocol based on rigorous methods in accordance with Cochrane and PRISMA 569 guidelines; the exhaustive search in all relevant databases; the independent screening of the titles, 570 abstracts and full-text of studies; the assessment of each included studies' risk of bias using 571 EPOC Cochrane guidelines; and the assessment of the quality of evidence for each individual 572 outcome using the GRADE methodology.

573 Our review also has limitations to consider. First, outcome measures varied widely across 574 studies. To address this issue, we conducted the meta-analysis using the SMD. Using the SMD 575 allowed us to standardize the results of studies to a uniform scale before pooling them. Review

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3 4	576	authors judged that using the SMD was the best option for this review, as it is the current practice
5 6 7 8 9 10 11 12 13 14	577	in the field of knowledge synthesis in medical education ^{6 60} .
	578	Second, there was high inconsistency among study results, which we can mostly attribute to
	579	differences in populations, AEE design, research methods, and outcomes. This resulted in
	580	sometimes widely differing estimates of effect. To partly address this issue, we used a random-
15 16	581	effects model for the meta-analysis, which assumes that the effects estimated in the studies are
17 18 19 20 21 22 23 24 25 26 27	582	different and follow a distribution ³⁰ . However, since a random-effects model awards more
	583	weight to smaller studies to learn about the distribution of effects, it could potentially exacerbate
	584	the effects of the bias in these studies ³⁰ .
	585	Finally, publication bias could not be assessed by the means of a funnel plot since there were less
27 28 29	586	than 10 studies included in the meta-analysis.
30 31 32	587	CONCLUSIONS
33 34 35	588	Adaptive e-learning has significant potential to increase the effectiveness and efficiency of
36 37 38 39 40 41 42	589	learning in health professionals and students. Through the different sub-domains of the adaptivity
	590	process (i.e. method, goals, timing, factors, types), AEEs can take into account the particularities
	591	inherent to each learner. This systematic review and meta-analysis underlines the potential of
43 44	592	AEEs for improving knowledge and skills in health professionals and students in comparison
45 46	593	with other educational interventions, such as nonadaptive e-learning environments and large-
47 48 49	594	group classroom learning, across a range of topics. However, evidence was either of low or very
50 51	595	low quality and heterogeneity was high across populations, interventions, comparators, and
52 53	596	outcomes. Thus, additional comparative studies assessing the efficacy of AEEs in health
54 55 56	597	professionals and students are needed to strengthen the quality of evidence .
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598 AUTHOR CONTRIBUTIONS

All review authors contributed to at least one aspect of each of the four criteria for authorshipdefined by the International Committee for Medical Journal Editors (ICJME).

G.F. contributed to the conception and design of the review, to the acquisition and analysis ofdata, and to the interpretation of results. Moreover, G.F. drafted the initial manuscript. S.C.

603 contributed to the conception and design of the review, and to the interpretation of results. M.-

604 A.M.-C. contributed to the conception and design of the review, to the acquisition of data, and

605 interpretation of results. T.M. contributed to the conception and design of the review, to the

606 acquisition of data, and to the interpretation of results. M.-F.D. contributed to the conception and

607 design of the review, to the acquisition of data, and to the interpretation of results. G.M.-D.

608 contributed to the conception and design of the review, and to the interpretation of results. J.C.

609 contributed to the interpretation of results. M.-P.G. contributed to the interpretation of results.

610 V.D. contributed to the interpretation of results.

All review authors contributed to manuscript writing, critically revised the manuscript, gave finalapproval, and agreed to be accountable for all aspects of work, ensuring integrity and accuracy.

613 DATA SHARING STATEMENT

614 All data relevant to the study are included in the article or uploaded as supplementary615 information.

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632 Figure 2. Risk of bias summary: review authors' judgements about each risk of bias item for each633 included study.

Figure 3. Risk of bias graph: review authors' judgements about each risk of bias item presented as
percentages across all included studies.

636 Figure 4. Forest plot representing the meta-analysis of the efficacy of adaptive e-learning versus637 other educational interventions in improving knowledge.

Figure 5. Forest plot representing the meta-analysis of the efficacy of adaptive e-learning versus
 other educational interventions in improving skills.
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45	Figure 2. Risk of bias summary: review author	rs'	iud	aen	nen	ts a	ihoi	it e	ach	risk of bias item for each included
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Figure 3. Risk of bias graph: review authors' judgements about each risk of bias item presented as percentages across all included studies.

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	Adaptiv	ve e-lear	ning	Other ed	ucational	interv.		Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Cook 2008	76.2	0.9	62	77.2	0.9	62	17.1%	-1.10 [-1.48, -0.73]	-
de Ruijter 2018	14.876	1.242	121	14.8447	1.1093	103	17.4%	0.03 [-0.24, 0.29]	+
Micheel 2017	7.861	1.691	282	7.461	1.691	317	17.6%	0.24 [0.08, 0.40]	*
Morente 2013	15.83	2.52	30	11.6	2.39	42	16.3%	1.71 [1.16, 2.26]	
Samulski 2017	5.83	2.43	18	4.47	2.47	18	15.7%	0.54 [-0.12, 1.21]	
Wong 2015	68.2	2.7	39	60.5	2.3	41	15.8%	3.05 [2.39, 3.70]	

Figure 4. Forest plot representing the meta-analysis of the efficacy of adaptive e-learning versus other educational interventions in improving knowledge.

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	Adapti	ve e-lear	ning	Other edu	icational i	nterv.	3	std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Hayes-Roth 2010	16.32	1.54	11	10.95	2.03	11	10.3%	2.87 [1.61, 4.12]	
Lee 2017	53	17	989	50	18	533	18.5%	0.17 [0.07, 0.28]	•
Romito 2016	67	4	12	53	6	12	11.1%	2.65 [1.51, 3.80]	
Thai 2015	69	16	27	57	23	27	16.1%	0.60 [0.05, 1.14]	
Van Es 2015	91.8	2.3	12	82.6	4.3	15	11.9%	2.51 [1.46, 3.55]	
Van Es 2016	71.7	18.73	15	58.2	20.2	18	14.8%	0.67 [-0.03, 1.38]	
Wong 2017	0.55	0.22	39	0.46	0.21	86	17.3%	0.42 [0.04, 0.80]	-
Total (95% CI)			1105			702	100.0%	1.19 [0.59, 1.79]	•
Heterogeneity: Tau ² =	0.50: Ch	$i^2 = 57.0$	0. df =	6 (P < 0.00)	$(001): I^2 =$	89%		-	

Figure 5. Forest plot representing the meta-analysis of the efficacy of adaptive e-learning versus other educational interventions in improving skills.

365x79mm (144 x 144 DPI)

PRISMA 2009 Checklist

Section/topic	#	Checklist item	Reported on page #
TITLE	-		
Title	1	Identify the report as a systematic review, meta-analysis, or both.	1
ABSTRACT	-		
Structured summary	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	4-5
INTRODUCTION	_		
Rationale	3	Describe the rationale for the review in the context of what is already known.	7-9
Objectives	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).	9-10
METHODS	_		
Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number.	10
Eligibility criteria	6	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.	10-11
Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.	11-12
Search	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.	Supplementary File 2
Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	11-12
Data collection process	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	11-12
Data items	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	12-13
Risk of bias in individual studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	13
Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	13-14
Synthesis of results	14	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., I ²) for each meta-analysis.	13-14

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PRISMA 2009 Checklist

5 Section/topic	#	Checklist item	Reported on page #
Risk of bias across studies	15	Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).	13
9 Additional analyses 10	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	14
RESULTS			
13 Study selection 14	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	15
15 16 17	18	For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and provide the citations.	18
8 Risk of bias within studies	19	Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	29-30
P Results of individual studies	20	For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.	32-33
2 Synthesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency.	32-33
Risk of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).	31
Additional analysis	23	Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).	N/A
27 DISCUSSION	-		
Summary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	32-33
31 Limitations 32	25	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of identified research, reporting bias).	39
Conclusions	26	Provide a general interpretation of the results in the context of other evidence, and implications for future research.	36-39
Funding	27	Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the systematic review.	41
40 41 <i>From:</i> Moher D, Liberati A, Tetzlaf 41 doi:10.1371/journal.pmed1000097	ff J, Altm	an DG, The PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. PLoS	Med 6(7): e1000097.
42 13		For more information, visit: www.prisma-statement.org.	
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Supplementary File 2 – Search Strategy for PubMed

PubMed - 17 avril 2017

- ((Adaptive[TIAB] OR individualized [TIAB] OR personalized[TIAB] OR Tailored[TIAB]) AND (elearning[TIAB] OR learning[TIAB] OR Instruction[TIAB] OR "web-based instruction"[TIAB] OR computer-based instruction[TIAB] OR computer-based tutoring[TIAB] OR education[TIAB] OR tutorial[TIAB] OR tutorials[TIAB])) OR Intelligent tutoring system[TIAB] OR Intelligent tutoring systems[TIAB]
- 2. "Computer-Assisted Instruction"[MH]
- 3. #1 OR #2
- 4. Health Personnel*[TIAB] OR Health professional*[TIAB] OR Health care profession*[TIAB] OR Healthcare profession*[TIAB] OR Medical student*[TIAB] OR Medical assistant*[TIAB] OR health worker*[TIAB] OR Audiologist*[TIAB] OR Chiropractor*[TIAB] OR Dentist[TIAB] OR Dentists[TIAB] OR Dietitian*[TIAB] OR Dermatolog*[TIAB] OR endocrinologist*[TIAB] OR Gastroenterolog*[TIAB]OR Gynecolog*[TIAB]OR Radiolog*[TIAB] OR Medical Staff[TIAB] OR Midwife*[TIAB] OR nutritionist*[TIAB] OR Nurse[TIAB] OR Nurses[TIAB] OR Optometrist*[TIAB] OR Occupational Therapist*[TIAB] OR Patholog*[TIAB] OR Paramedic[TIAB] OR Paediatric[TIAB] OR pediatrician*[TIAB] OR Paediatrician*[TIAB] OR pediatrist*[TIAB] OR pediatric[TIAB] OR Pharmacist*[TIAB] OR Pharmaconomist*[TIAB] OR Pharmacologist*[TIAB] OR Pharmacy technician*[TIAB] OR Phlebotomist*[TIAB] OR Physician OR Podiatrist*[TIAB] OR Psychologist*[TIAB] OR Psychotherapist*[TIAB] OR psychiatrist*[TIAB] OR Physical therapist*[TIAB] OR physiotherapist*[TIAB] OR Respiratory therapist*[TIAB] OR Surgeon*[TIAB] OR Clinician*[TIAB] OR Cardiologist*[TIAB] OR Emergency medical technician*[TIAB] OR emergency doctor*[TIAB] OR emergentologist*[TIAB] OR clinical officer*[TIAB] OR Community health worker*[TIAB] OR Radiographer*[TIAB] OR Surgical technologist*[TIAB] OR Radiotherapist*[TIAB] OR Anesthetist*[TIAB] OR Resident[TIAB] OR residents[TIAB]
- "Health Personnel"[MH] OR "Students, Premedical"[MH] OR "Students, Medical"[MH] OR "Students, Nursing"[Mesh]
- "Education, Premedical"[MH] OR "Education, Medical"[MH] OR "Education, Nursing"[MH] OR "Education, Pharmacy"[MH] OR "Education, Public Health Professional"[MH] OR "Clinical Clerkship"[MH]
- 7. #4 OR #5 OR 6
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- 10. #8 OR 9
- 11. #3 AND #7 AND #10
- 12. (english[LA] OR french[LA]) AND 2005:2017[DP]
- 13. #11 AND #12

Résultats: 4 375

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Supplementary File 3 – Summary of the quality of evidence

Certainty assessment						№ of patients		Effect					
Nº of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	adaptive and intelligent e- learning environments	other educational interventions	Relative (95% Cl)	Absolute (95% Cl)	Certainty	Importance	
Knowledge	Knowledge												
6	randomised trials	serious ^a	serious ^b	not serious	serious °	none	552	583	-	SMD 0.7 SD higher (0.08 lower to 1.49 higher)		IMPORTANT	
Competence	Competence												
7	randomised trials	serious ^a	not serious	not serious	serious °	none	1105	702	-	SMD 1.19 SD higher (0.59 higher to 1.79 higher)		CRITICAL	
CI: Confidence	CI: Confidence interval; SMD: Standardised mean difference												
Explanations													
a. Most studies have unclear or high risk of bias with regard to random sequence generation and allocation concealment. The risk of bias for similarity of baseline measurements was unclear for some studies. Thus, groups in these studies could be disproportionate and the distribution may not be normal since sample size is generally small.													
				For pee	er review on	ly - http://bmjopen	.bmj.com/site	/about/guidel	ines.xhtml				