Meta-analysis of online smoking cessation interventions

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Shahab, L. & McEwen, A (2009). Online support for smoking cessation – a systematic review of the literature. *Addiction*, **104**, 1792-1804.

Online support for smoking cessation: a systematic review of the literature

Short title: Meta-analysis of online smoking cessation interventions

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Word Count: 4342 (excluding Abstract=221)

Pages: 35

Figures: 4

Tables: 3

This review was funded by the Department of Health, England. Lion Shahab has received an honorarium for a talk and travel expenses form a pharmaceutical company making smoking cessation products. Andy McEwen has received travel funding, honorariums and consultancy payments from manufacturers of smoking cessation products (Pfizer Ltd, Novartis and GSK Consumer Healthcare Ltd).

Online support for smoking cessation: a systematic review of the literature

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ABSTRACT

Aim: To examine the efficacy and acceptability of online, interactive interventions for smoking cessation and to identify treatment effect moderators and mediators.

Methods: A systematic review and meta-analysis of the literature (1990-2008) was conducted, finding eleven relevant randomised controlled trials. Data were extracted and risk ratios and risk differences estimated with a random effects model.

Results: There was no evidence of publication bias. Included trials were of variable methodological quality. Web-based, tailored, interactive smoking cessation interventions were effective compared with untailored booklet or email interventions (RR 1.8; 95%CI 1.4-2.3) increasing 6-months abstinence by 17% (95%CI 12-21%). No overall effect of interactive compared with static Web-based interventions was detected but there was significant heterogeneity with one study obtaining a clear effect and another failing to find one. Few moderating or mediating factors were evaluated in studies and those that were had little effect. Pooled results suggest that only interventions aimed at smokers motivated to quit were effective (RR 1.3, 95%CI 1.0-1.7). Fully automated interventions increased smoking cessation rates (RR 1.4, 95%CI 1.0-2.0) but evidence was less clear-cut for non-automated interventions. Overall, the Web-based interventions evaluated were considered to be acceptable and user satisfaction was generally high.

Conclusion: Interactive, Web-based interventions for smoking cessation can be effective in aiding cessation. More research is needed to evaluate the relative efficacy of interactive Web-based interventions compared with static websites.

Key words: online health promotion; Web-based intervention; smoking cessation; metaanalysis; systematic review

INTRODUCTION

Smoking is one of the most significant factors contributing to low life expectancy, health inequalities and ill health worldwide [1]. Smoking prevalence has fallen considerably, especially in developed countries, and yet less than half of smokers ever achieve long-term abstinence despite 70-80% wanting to quit and one-third having made at least three serious quit attempts [2]. This underlines the need for the development of new methods that prevent people from starting to smoke, motivate smokers to quit smoking and sustain long-term cessation.

Effective smoking cessation interventions exist [3]. However, the most effective interventions (such as face-to-face counselling) also have the lowest reach and interventions with the widest reach (e.g. mass media campaigns) have low efficacy [4]. One way forward is to either increase the reach of intensive, low-reach interventions or the efficacy of less intensive, high-reach interventions [5]. Tailoring of information has been shown to increase effectiveness of low intensity interventions such as self-help materials [6] but these do not allow smokers to interact as they can in face-to-face interventions. With advances in computer-based communication technologies, the ladvantages of wide-reaching interventions through incorporating interactive responses [7] and thus adopting some of the features of face-to-face contact [8]. Moreover, Internet-based interventions can exploit economies of scale and are therefore potentially very cost effective [9].

Currently, around 75% of the population in industrialised countries and around 10% in the rest of the world have access to the Internet [10;11], and these proportions are likely to rise further. Online support for smoking cessation therefore represents a viable

resource for smokers wanting to stop, especially because it can be accessed at any time from anywhere where there is an Internet connection [12]. However, studies of online smoking cessation resources find great variability in quality and acceptability: up to 80% of reviewed websites provided no information on key components endorsed by smoking cessation clinical guidelines [12]. Consequently, many users report being unsatisfied with Web-based smoking cessation interventions, finding them confusing and unhelpful [13]. Moreover, despite an increase in the number of online interventions, relatively little is known as to how and why these interventions work and which factors influence their effectiveness [14].

Several reviews have evaluated the use of Internet-based interventions to change health behaviours, including physical exercise, weight loss, disease management and prevention [see 15;16;17;18;19]. While treatment efficacy differs with the health behaviour that is targeted, these reviews generally support the usefulness of Web-based interventions for health promotion. Two reviews that focused exclusively on smoking cessation interventions [20;21] concluded that computer-based, including Web-based, interventions have the potential to increase smoking cessation rates. However, both reviews included interventions that, although computer-based, were not interactive or delivered online.

The delivery and content of websites is not regulated [22] and it is easy to imagine how variability across websites could make generalisations about this type of intervention problematic. However, it is possible to assess the potential for this kind of intervention to provide smoking cessation support by examining the efficacy found in rigorous evaluations carried out to date. To our knowledge, this is the first meta-analysis to evaluate the efficacy of interactive, online smoking cessation interventions compared

with either low-efficacy interventions with high potential reach (e.g. self-help materials such as booklets) or with high-efficacy interventions of low potential reach (e.g. face-to-face counselling). This review also includes studies not covered in other published reviews. Moreover, we aimed to clarify possible mechanisms of action by identifying potential moderators and mediators (i.e. specific intervention components) of the treatment effect.

METHODS

Study selection

To be included in this review articles had to be in English, published since 1990, have a randomized controlled trial (RCT) design with at least one month follow-up assessing smoking cessation and participants had to be current smokers at enrolment. In order to differentiate the potential of Internet-based interventions from traditional self-help material, only interventions that made use of the interactive nature of the Internet were included. Multiple follow-up studies that used data from the same cohort of participants were excluded to avoid bias in effect estimates [23]. Studies providing insufficient detail/data and those focusing on secondary outcomes (such as training health professionals involved in providing smoking cessation interventions) were also excluded.

Data source and search strategies

Relevant literature was sought by searches of computerised databases carried out in December 2008. The electronic resources used were PubMed (1990 – 2008/12), PsycINFO (1990 - 2008/12), CINHAL Plus (1990 - 2008), EconLit (1990 - 2008/11), ISI Web of Science[®] (SCI, SSCI, AHCI; 1990 - 2008) and CENTRAL (Cochrane Registry of Controlled Trials, 1990 – 2008). Medline was searched with relevant MeSH terms ((("Online Systems" OR "Internet") AND "Smoking Cessation") AND "Randomized

Controlled Trial"). All other databases were searched with free text terms reflecting inclusion criteria: ((("Internet" OR "Online" OR "Web") AND "Smoking cessation") AND ("RCT" OR "Randomi\$ed controlled trial" OR "Random* trial")). This yielded 82 studies, reduced to 14 studies when reading the abstracts and nine suitable studies after scrutinizing the full-text of articles. Further free text and bibliography searches revealed two additional articles that fulfilled selection criteria, bringing the total of included studies to eleven [24-34]. Egger's regression intercept as well as the Begg and Madzumdar rank correlation coefficient, which are used to quantify publication bias [35], were non-significant, suggesting that no file drawer effect was present.

Data extraction and analysis

Data were extracted manually and recorded on a data extraction sheet to minimise bias when coding and rating studies [36]. Results were analysed with RevMan [37] and presented as rate ratios and, where appropriate, risk difference was calculated. The extent of heterogeneity between study outcomes was assessed using the I² statistic, which is based on Cochrane's *Q* but is more robust and thus suitable for smaller meta-analyses [38]. Where appropriate, studies were pooled with the Mantel-Haenzel method using random effect rather than fixed effect models to maximise accuracy of parameter estimation [39].

Study Quality

Attrition rates tend to be high in Internet-based interventions [40]. Indeed, attrition in studies varied widely (Table 1), suggesting that some of the results would be affected by attrition bias. All studies (except [32]) therefore used an intention-to-treat analysis. The methods of recruitment and the populations sampled were reasonably homogeneous (Table 2). However, little information was given about participants included and excluded

from studies, making it impossible to evaluate how representative the included samples were. Studies also provided little detail on intervention characteristics and whether outcome assessors were blinded.

Most studies (N=10) used well-validated questionnaires to measure constructs of interest. Yet, validation of smoking cessation outcome was limited: only two studies objectively verified smoking status, either in a sub-sample [24] or in the total sample [30]; the remainder relied on self-report. There were inconsistencies in the definition of abstinence between studies; 7 or 30-day point prevalence and continuous abstinence were used in different studies.

Table 1 about here

Only one study [24] detected baseline differences between groups but information on socio-demographic and smoking characteristics was lacking for a number of study samples (Table 3). Treatment and control group content were often not well matched (Tables 1, 2 and 3). This imbalance between control and intervention conditions makes it harder to evaluate whether differences between groups were due to the intervention or confounding variables such as direct contact time.

Details on statistical procedures was often missing (Table 1); just two studies [24;31] reported use of power calculations to estimate the required sample size before recruitment, and one of these postulated an optimistic group difference of 12% [24]. A realistic treatment effect size of 2-5% (i.e. a small effect [Cohen's w=.1]) would require a minimum sample of 393 participants per group to detect this difference at a significance level of 5% with a power of 80%. Only three studies in this review [28;31;33] reached

this number of participants, while two studies [30;32] were in danger of Type II error because they were severely underpowered (N<100 per group).

RESULTS

Study Description

All eleven reviewed studies were published since 2005, included a total of 15,511 participants and most (N=8) were carried out in the USA (Table 2). Participants were largely recruited from the general population; with the exception of two studies [24;30] that targeted college students and adolescents. While most trials required participants to intend to stop smoking to be eligible, four studies did not explicitly state or require that smokers had to be willing to stop (Table 2).

Table 2 about here

Studies included mainly white, female smokers with at least high school education of a wide age range (Table 3). There was great variability in cigarette consumption as some studies [notably 24] included non-daily smokers; however, smokers in most studies consumed at least ten cigarettes a day.

The majority of RCTs compared an interactive Internet-based intervention with either a minimal control condition (booklet or a static website containing information derived from standard smoking cessation material) or with a waiting-list control (Table 2). Two studies compared an interactive, Internet-based intervention with face-to-face counselling [30;32] and two RCTs [27;32-this study had two control conditions] evaluated the efficacy of an Internet intervention as an adjuvant to a behavioural intervention compared with the behavioural intervention alone. Two further studies contrasted an

interactive Web-based control intervention with an interactive online intervention that was 'enhanced' by either including more targeted, comprehensive material [28] or by having additional treatment components [29] (Table 2).

As a minimum, all interventions included baseline and follow-up assessments to tailor provision of information, structure quit plans and deliver relapse prevention advice. Tailoring (the iterative and ipsative use of input information to structure output) is only one way to make use of the inherent interactive nature of the Internet; most studies (N=8) also employed emails and/or text messages to contact participants periodically and to provide support material. In addition, they supplied links to other websites with further information and used quizzes to consolidate disseminated information. Four studies used reminder phone calls or emails if participants had not been in contact for a predetermined period of time (Table 2).

Table 3 about here

Six interventions were fully automated; the remainder contained some opportunity for smokers to interact with 'peer coaches' or 'experts' to ask for advice, as well as allowing participants to interact with peers via chat rooms (Table 2). Three interventions [30;31;34] featured role models, mostly through the use of videos that were matched to smokers' characteristics with the aim of increasing motivation and self-efficacy through social modelling; Brendryen et al. [25;26] provided an automated craving line that participants could call if they felt the urge to smoke. Two studies [25;27] provided concomitant pharmacotherapy while one study [33] only included participants that had purchased NRT over the counter.

The majority of intervention programs were loosely based on evidence-based practice guidelines [41], social cognitive theory [42], emotional writing paradigm [43] and the stages of change model [44]. Overall, the length of treatment varied considerably from 10 weeks [33] to one year [25], as did follow-up duration (Table 2 and 3).

There was variability in the control conditions provided (Table 3). Only four studies compared the Web-based intervention with an equivalently complex control intervention [28-30;32]. A number of studies attempted to mimic the intervention in the control condition by providing a static website containing general information about smoking and smoking cessation based on standard smoking cessation manuals and booklets (Table 2). Two studies offered only minimal [24] or no control intervention [34].

Study Outcomes

Smoking cessation

At the final follow-up point, five studies [24-26;33;34] found an overall positive effect of the intervention on smoking cessation rates. Three further studies found an effect in subgroup analyses [27;30;31] and three studies did not detect a positive effect of the intervention [28;29;32]. As study methodology and intervention content were diverse, we did not pool study results overall. However, in order to present some meaningful data and quantify individual results, we attempted to decrease heterogeneity and increase reliability of findings across studies by analysing and pooling study results according to common methodological criteria.

A random effects model of studies that compared interactive, tailored Internet smoking cessation interventions with minimal control conditions (i.e. booklet/email/static website/waiting list) found a significant treatment effect (RR 1.5; 95% CI 1.2-1.9,

p<0.001). However, there was significant heterogeneity in effect size. The increase in quit rates was larger when interactive, online smoking cessation interventions were compared with a booklet/email than with a static website control condition (Figure 1, Sub-category 2 and 3).

Figure 1 about here

Separate analysis by control condition confirmed a significant non-heterogeneous treatment effect for comparisons with booklet or email control condition (RR 1.8; 95%CI 1.4-2.3, p<0.001, Sub-category 2). However, the effect in studies with a static website control was non-significant and heterogeneous (Sub-category 3) as this included a large trial [31] that did not detect a treatment effect. This study found differences in access to different websites used in the intervention condition; participants randomized to the most highly accessed intervention websites had significantly higher point-prevalence (but not continuous) abstinence rates than those randomized to the other websites. In a sub-analysis that included only participants allocated to the heavily utilized intervention websites, we found that heterogeneity was no longer significant and results supported the treatment effect of tailored, interactive smoking cessation websites (RR 1.2, 95%CI 1.1-1.4, p<0.001, Sub-category 3a).

Figure 2 about here

As about half of people that are continuously abstinent at six months achieve long-term smoking cessation, these data can be used to estimate permanent abstinence rates [45]. We therefore pooled the results of studies that provided half-year follow-up data on the percentage difference in success rates [24-26]. There was little heterogeneity; a random

effects model suggests that interactive, online interventions increased abstinence by 17% (95%CI 12-21%) effectively doubling smoking cessation rates compared with minimal control conditions (RR 1.9, 95%CI 1.9-2.6, p<0.001; Figure 3). However, it should be noted that these data are based on point prevalence not continuous abstinence rates, which is likely to result in greater relapse rates.

Two studies compared an interactive Internet intervention with either face-to-face counselling [30] or face-to-face motivational interviewing [32]. Neither study found evidence that the Internet intervention increased smoking cessation rates compared with the control intervention (RR 0.6, 95%CI 0.3-1.1; Figure 3, Sub-category 1). Similarly there was no evidence that Web-based interventions increased abstinence in smokers also using face-to-face interventions (RR 1.3, 95%CI 0.8-2.1; Figure 3; Sub-category 2). However, the aggregate sample size in both these cases was small so the power to detect an effect would be very limited.

Figure 3 about here

A further two studies evaluated the impact of 'enhanced' Internet smoking cessation interventions on abstinence rates. One compared an interactive Internet intervention that incorporated a mood management component with the same intervention lacking this component in two different settings: one was delivered in English and one in Spanish to appeal to Hispanic smokers in the US [29]. Another study compared an interactive online intervention, including comprehensive, targeted components to increase smoking cessation, with a control that contained less explicit support for smoking cessation but aimed to increase physical activity as a way to stop smoking. Neither study found an effect of the enhanced intervention (RR 0.9, 95%CI 0.6-1.2; Figure 4); in fact when

results were analysed separately according to language modality in the first trial, the addition of the mood management component depressed smoking cessation rates at 12 months for users of the English version of the intervention ($X^2(1)=4.4$, p=0.04).

Figure 4 about here

As study methodologies were heterogeneous, we did not pool results according to follow-up point. However, individual studies that included several follow-ups provide some insight into the length of the treatment effect. Studies that did not find an initial treatment effect did not detect a late effect [27-30], whereas trials that reported a significant effect found this persisted throughout subsequent follow-ups [24-26;33], up to 12 months following some interventions.

Effect moderators and mediators

Whilst a number of baseline variables commonly associated with smokers' chances of succeeding in quitting were identified (e.g. nicotine addiction [24], depression [29]), no interactions between intervention and baseline characteristics (including type of Internet access – dial-up vs. broadband) were observed when this was assessed.

Intention to stop smoking is an important putative moderator of intervention effectiveness. Pooled results from trials that did not require participants to intend to stop smoking in the near future (N=4, Table 2) indicate no intervention effect (RR 1.2, 95%CI 0.8-1.9) but outcomes were heterogeneous (I^2 =77.6%; $\chi^2(3)$ =13.4, p=0.004). In contrast, while equally heterogeneous (I^2 =70.3%; $\chi^2(6)$ =20.2, p=0.003), the pooled results from trials recruiting smokers motivated to quit (N=7) show a treatment effect (RR 1.3, 95%CI 1.0-1.7, p=0.04).

As all interventions tested contained multiple components, it is important to try to determine which of these influenced outcomes. When pooling studies according to the length of treatment provided (see Table 3), there was no clear association with increased cessation rates. The pooled results of studies that were fully automated (N=6, Table 2) showed a treatment effect (RR 1.4, 95%Cl 1.0-2.0, p=0.03) but results were heterogeneous (l^2 =73.8%; χ^2 (5)=19.1, p=0.002). In contrast, the pooled result of the five non-automated studies was not significant (RR 1.2, 95%Cl 0.9-1.6) but also heterogeneous (l^2 =62.7%; χ^2 (4)=10.7, p=0.03).

Two studies that reported an increase in NRT use found that the effect of an interactive, online smoking cessation intervention was not mediated by usage rate of pharmacotherapy [25;33]. This was confirmed by two trials that found the same intervention to be effective with and without the provision of NRT [25;26]. These studies also observed that self-efficacy, at least partially, mediated the impact of the intervention. Lastly, in trials where this was assessed, a greater frequency of use of the provided Internet intervention was associated with higher abstinence rates. However, it was not possible to evaluate further moderators and mediators as interventions were both too diverse and did not provide sufficient detail to carry out further subanalyses.

Website utilization and satisfaction

As shown in Table 4, loss to follow-up and drop-out from the Web-based smoking cessation interventions was high, which is mirrored when looking at the usage of interactive websites. The proportion of users allocated to the treatment condition that logged on at least once was reasonably high (e.g. 70.2% [34]) but the use of the websites quickly tapered over time [27;28;30]. Brendryen *et al* [25] contacted participants

that had left the programme to identify reasons for their decision to discontinue and reported that this was mostly due to stressful life events or the intrusiveness of the Webbased intervention.

To assess participants' views of the content of online interventions, Japunitch and colleagues [27] investigated the use of different components of their intervention and found that support tools (discussion groups and chat rooms, including an 'ask the expert' service) were the most popular module of their programme, while information tools (providing educational material about smoking cessation) were least popular. However, Swartz *et al* [34] did not find any differences in the utilisation of various modules of their online smoking cessation programme. Satisfaction with treatment programmes for smoking cessation was generally very high at around 90% [25;26;33] and tailored intervention fared significantly better than non-tailored static Internet interventions where this was compared [33].

DISCUSSION

Any meta-analysis, such as this, that considers a very diverse and heterogeneous set of interventions will necessarily be limited in the generalisability of its conclusions. Moreover, given the lack of detailed description of interventions further complicates the analysis and evaluation of mediating and moderating effects. However, using stringent inclusion criteria and classifying interventions according to methodological similarities provides at least some measure of control and insight into the relative efficacy of the interventions. Studies evaluating tailored, interactive Internet interventions compared with untailored written or Web-based material detected a significant treatment effect across different follow-up times and treatment schedules; online interventions tested so far increased abstinence rates by an estimated 17 percentage points (95%CI 12-21%) at

six months follow up. Given that unaided quit attempts result in abstinence rates of less than 5% at six months [46], Internet interventions have the potential to more than double long-term abstinence rates. Moreover, the treatment effect of Web-based smoking cessation interventions appears relatively stable over time as differences in cessation rates between control and treatment groups did not dissipate at subsequent follow-ups.

These results compare favourably with findings from meta-analyses evaluating other smoking cessation interventions of similar potential reach, which typically increase abstinence rates by between one and three percent [47-49]. However, whilst RCTs that directly compared Internet interventions with self-help materials found a clear treatment effect, no direct comparisons with telephone counselling or brief advice were available. Abstinence rates from Internet interventions were comparable to those usually obtained from more intensive, low reach interventions such as face-to-face individual or group counselling [50;51]. Indeed, there were no differences in cessation rates in trials that directly compared Internet-based interventions with face-to-face counselling but these studies had small samples. Adding an online smoking cessation intervention to behavioural treatment or enhancing Internet interventions did not confer any additional benefits. It therefore appears that online Internet interventions for smoking cessation have the potential to combine the wide reach of less intensive interventions with the greater efficacy of face-to-face counselling. Findings from a study evaluating the efficacy of online interactive smoking cessation treatment for smokeless tobacco users indicate that such interventions can also be effective for other forms of tobacco use and different groups of smokers [52]. This is in agreement with the overall efficacy of interventions that were included in this review, which targeting diverse populations including adolescents.

Similar to earlier reviews [15;21] we found considerable heterogeneity in the theoretical basis of interventions and no common rationale for the inclusion of various treatment components. As previously reported [6], tailoring of information was effective. Comparison with static websites suggests that an interactive user interface boosts the treatment effect although it is unclear what form of interaction is required to confer this benefit. While it has been speculated that chat forums could aid smoking cessation through the provision of additional social support [53;54], it appears that completely automated interventions may be more effective. There was no evidence that increasing treatment complexity improved cessation rates but this could reflect a ceiling effect as all reviewed interventions were already highly complex. However, reducing the complexity of interventions can lead to a reduction in treatment efficacy [55], thus striking the right balance between user involvement and treatment intricacy is important for determining the impact of an intervention.

Few treatment moderators and mediators were explicitly examined in studies. While insufficient description precluded an in-depth analysis of moderator and mediator effects, stratifying studies according to common methodological characteristics allowed some delineation of possible moderating effects. There was generally no impact of socio-demographic characteristics but motivation to quit appeared to influence the success of interventions. This lack of socio-demographic moderators may be surprising; however, given that all interventions attempted to match the provided treatment according to target population characteristics, one would not expect a differential outcome between various subgroups. This is in agreement with previous research that found few socio-demographic moderating variables and perceived relevance of the intervention as the only mediator of successful cessation [56].

Although there is considerable variability in the quality and usability of existing online interventions [13], use of, and reported satisfaction with, Web-based smoking cessation interventions was generally high. Nonetheless, usability testing by potential users of online programmes, rather than just by smoking cessation and Web development experts, is likely to further improve existing online interventions [57].

Internet-based health interventions provide obvious methodological challenges [58;59] that need to be acknowledged. First, biochemical outcome validation is impractical in Internet-based interventions as they span large geographic areas and due to little contact between treatment provider and client generally acknowledged not to be necessary [60]. Interestingly, studies that did validate abstinence did not report a difference in deception rate between intervention and control conditions [24;30]. Second, the use of the Internet as a medium for delivering interventions may bias against people with low computer literacy; i.e. introduce a 'digital divide' [58]. However, as the socio-demographic correlates of the digital divide have become blurred [61], Web-based interventions have increasingly shown their potential to involve users commonly excluded from research [62].

Third, the representativeness of trial participants needs to be established as substantial differences exist between smokers that are approached to participate and those that enrol with Internet smoking cessation trials [63]. Only two studies in this review compared characteristics of study participants with those of smokers who were approached or with the general population [30;31]. Another issue is the ability of Internet based interventions to attract and retain participants in research trials [64]; underlined by the fact that most of the included trials were underpowered due to a small sample and high attrition. This is likely to have undermined what can reliably be said about the

magnitude of observed effects. Lastly, the relatively small number of included studies and variable methodological quality somewhat limits the generalisability of the reported findings. As with many behaviour change interventions [65], most studies provided inadequate detail on treatment characteristics and more needs to be done to improve reporting. In particular, most studies provided little information about structural factors that may be of importance to website users such as the navigational, organisational and graphic interface or pacing, the impact of which could therefore not be examined in this review.

With these caveats in mind, it appears that online interventions for smoking cessation are acceptable to users and of superior efficacy to other wide-reach interventions. Current, limited evidence also suggests that online interventions may have similar efficacy to face-to-face interventions. The impact of Web-based smoking cessation interventions on quit rates seems to be relatively long-lasting and appears to be mediated by appropriate tailoring of information and personalising the experience through interactive online components. Web-based Internet interventions seem particularly effective for smokers intending to quit but few other differentiating factors were observed. Some methodological concerns exist. It would be desirable for future trials of online smoking cessation interventions to provide adequate description, to evaluate the actual reach of interventions and the representativeness of included participants and to clearly delineate theoretical underpinnings of included components. More research is required to confirm the relative efficacy of interactive online interventions compared with static websites and face-to-face counselling.

Acknowledgements

This review was funded by the Department of Health, England. Lion Shahab has received an honorarium for a talk and travel expenses form a pharmaceutical company making smoking cessation products. Andy McEwen has received travel funding, honorariums and consultancy payments from manufacturers of smoking cessation products (Pfizer Ltd, Novartis and GSK Consumer Healthcare Ltd). We would like to thank Robert West for help in preparing this manuscript.

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Tables

Table 1: Study methodology

Reference	True Rando- misation	Matched group characteristi cs	Matched treatme nt	Standardise d Procedures	Train ed Staff	Measure & outcome validation	Blinded assesso r	Attritio n rate⁺(%)	ITT analys is	Power & statistical analyisis
An (2008) [24]	Yes	Intervention group older, no differences on other measures	Minimal	Yes	Not know n	Measures validated but abstinence only partly confirmed with biomarker (CO); 30 day abstinence	Not known	C: 8.8 I: 7.0	Yes	β=0.85 to detect 12% difference; little detail on stats: mostly logistic regression; stat assumptions not tested
Brendryen (2008a) [25]	Yes	No difference on small set of measures	Somewh at	Yes	N/A	Measures validated except self-efficacy measure; abstinence not biochemically validated; 7 day point prevalence	Not known	C: 15.6 I: 12.2	Yes	Powered but not specified; detailed stats: χ^2 , t-test, logistic regression; stat assumptions not tested
Brendryen (2008b) [26]	Yes	No differences on extensive set of measures	Somewh at	Yes	N/A	Measures validated except self-efficacy measure; abstinence not biochemically validated; Repeated 7-day point abstinence	Not known	C: 26.0 I: 18.1	Yes	Power unclear; detailed stats: χ^2 , t- test, logistic regression; stat assumptions not tested
Japuntich (2006) [27]	Unclear	No test of differences	Minimal	Yes	Yes	Measures validated; abstinence validation unclear; 7 day point prevalence	Not known	C: 20.8 I: 19.3	Yes	Power unclear; little detail on stats: mostly logistic regression

¹ The use of alternation, case record numbers, birth dates or week days is considered to be inadequate for randomisation (Verhagen, de Vet, de Bie, Kessels *et al.*, 1998); ⁺ Attrition rate refers to cumulative loss to follow-up at last contact point; C-Control condition, I-Intervention.

Reference	True Rando- misation ¹	Matched group characteristi cs	Matched treatme nt	Standardise d Procedures	Train ed Staff	Measure & outcome validation	Blinded assesso r	Attritio n rate⁺(%)	ITT analys is	Power & statistical analyisis
МсКау (2008) [28]	Unclear	No differences on extensive set of measures	Yes	Yes	Not know n	Measures validated except self-efficacy, and depression abstinence not biochemically validated; 7 day point prevalence	Not known	C: 72.6 I: 72.9	Yes	Powered but not specified; detailed stats: logistic regression, Kaplan- Meier survival analysis; stat assumptions not tested
Munoz (2006) [29]	Unclear	No test of differences	Yes	Yes	N/A	Measures validated, abstinence not biochemically confirmed; 7 day point prevalence	Not known	C:52.6 I:50.9	Yes	Power unclear; little detail on stats: χ^2 test
Patten (2006) [30]	Unclear	No differences on extensive set of measures	Yes	Yes	Yes	Measures validated; cessation confirmed by CO; 30 day abstinence	Not known	C: 48 I: 41	Yes	Power unclear; detailed stats: χ^2 , Fisher's exact test, rank sum test, GEE
Pike (2007) [31]	Unclear	No differences between groups on unknown set of measures	Minimal	Yes	Not know n	No validation of measures provided; abstinence not biochemically confirmed; 7 day point prevalence	Not known	46 – similar in both groups	Yes	β =0.8 to detect 3% difference; little detail on stats: χ^2 and t-test, stat assumptions not tested
Prochaska (2008) [32]	Unclear	Cannot be estimated	Yes	Yes	Yes	Measures, validated, abstinence not biochemically confirmed; Point prevalence	Not known	25.7 – not provide d by group	No	Power unclear; little detail on stats: χ ² and Levy test for pair-wise proportions

Reference	True Rando- misation	Matched group characteristi cs	Matched treatme nt	Standardise d Procedures	Train ed Staff	Measure & outcome validation	Blinded assesso r	Attritio n rate ⁺ (%)	ITT analys is	Power & statistical analyisis
Strecher (2005) [33]	Unclear	No differences between groups on unknown set of measures	Somewh at	Yes	N/A	Measure validation not known, abstinence not biochemically confirmed; 10 week abstinence	Not known	C: 78.9 l: 77.6	Yes	Power unclear; little detail on stats: logistic regression, stat assumptions not tested
Swartz (2006) [34]	Yes	No differences between groups on extensive set of measures	Not at all	Yes	N/A	Measure validation unclear, abstinence not biochemically confirmed; 7 day point prevalence	Not known	C: 38.1 I: 49.1	Yes	Power unclear; detailed stats: χ^2 and logistic regression

Table 2: Study summary

Reference	Sample and setting	Recruitment	Intervention	Follow-up	Design	Main Findings
An (2008)	517 college smokers at University of MN; USA; Not required to intend to stop smoking	College Internet health screening and follow-up email	RealU: Website based on social cognitive and problem behaviour therapy; logs health/lifestyle habits, has interactive quiz & feedback, links to lifestyle magazine, provision of weekly, tailored emails from peer coaches; provision of financial incentive for participation	8, 20, 30 weeks	RCT-2 groups: (i) Control condition (email with link to online health and academic resources); (ii) RealU intervention	Participants in (ii) more likely to report having stopped smoking for a month at follow-up
Brendryen (2008a)	396 smokers recruited from general population; Norway; Required to intend to stop smoking	Banner advertisement in Internet newspaper	Happy Ending: Daily email to direct to webpage; pre-recorded daily audio message (log-on call) and up to three text messages, proactive log-off call in evening to assess smoking and provide relapse prevention information when necessary, plus directs to craving helpline; fully automated tunnel design; participants received reminders	1, 3, 6, 12 months	RCT-2 groups: (i) Booklet intervention (booklet with information, quit calendar, telephone number & links to online resources + NRT sample); (ii) Happy Ending intervention + NRT sample	Participants in (ii) significantly more likely to have stopped smoking and to adhere to NRT
Brendryen (2008b)	290 smokers recruited from general population; Norway; Required to intend to stop smoking	Banner advertisement in Internet newspaper	Happy Ending: Daily email to direct to webpage; pre-recorded daily audio message (log-on call) and up to three text messages, proactive log-off call in evening to assess smoking and provide relapse prevention information when necessary, plus directs to craving helpline; fully automated tunnel design; participants received reminders	1, 3, 6, 12 months	RCT-2 groups: (i) Booklet intervention (booklet with information, quit calendar, telephone number & links to online resources); (ii) Happy Ending intervention	Participants in (ii) significantly more likely to have stopped smoking and to have higher levels of coping planning and pre-cessation self- efficacy
Japuntich (2006)	284 smokers recruited from general population in WI; USA; Required to intend to stop smoking	Billboards, posters, fliers, TV advertisement, press releases	CHESS SCRP: Online system with log in leading to tailored feedback, information provision, support centre providing Web- based cognitive-behavioural techniques (chat room/ask the expert) and folder to carry out emotional writing paradigm/keep diary; participants received reminders	3, 6 months	RCT-2 groups: (i) Standard intervention: 9 weeks buproprion, 3 brief counselling sessions and 5 follow-up visits; (ii) as (i) plus CHESS SCRP	No difference in cessation rates between (i) and (ii) but actual use of online system predicted cessation

Reference	Sample and setting	Recruitment	Intervention	Follow-up	Design	Main Findings
МсКау (2008)	2318 smokers recruited from general population in USA; Required to intend to stop smoking smoking	Banner advertisement on Internet search engines	QSN: Tailored (tunnel design), interactive webpage providing educational information, cessation preparation, personal quit plan and date, behavioural tasks, thought and mood management, peer support (chat room), expert advice and prompts	3, 6 months	RCT-2 groups: (i) Active Lives – online personalised fitness programme to encourage cessation comprising multi-step plan with motivational component, action plan with tracking, online resources and peer support (chat room); (ii) QSN	No differences in abstinence rates between (i) and (ii) possibly due to low engagement with online programmes
Munoz (2006)	568 English or Spanish speaking smokers recruited from general population in USA; Required to intend to stop smoking	Press release and Internet	Guia+ITEMS+MM: Web-based intervention of brochure (Guia) providing standard cessation information, tailored advice; itemised encouraging reminder emails (ITEMS); online mood management course (MM) providing instructions about monitoring feelings and relaxation; fully automated; financial incentive	1, 3, 6, 12 months	RCT-2 groups: (i) Guia+ITEMS; (ii) Guia+ITEMS+MM Carried out either as integrated in Spanish or English website	In separate analysis for Spanish and English websites studies, lower abstinence rates in (ii) than (i) for latter website
Patten (2006)	139 adolescent smokers from MN, WI, CT; USA; Not required to intend to stop smoking	TV, radio newspaper advertisement, fliers in schools and health clinics	SOS: Web-based with 38 components including interactive services (quizzes, tailored quit plans and notes, Q&A email service, online discussion support group), information services and videos (showing personal stories of quitters); financial incentive	8, 12, 24 (36) weeks	RCT-2 groups: (i) Brief office intervention (4 sessions of motivational interviewing/information & homework); (ii) SOS intervention	No difference in cessation rates between (i) and (ii) but decrease in days smoked in (i)
Pike (2007)	6451 smokers recruited from general population; USA; Not required to intend to stop smoking	Advertisement on American Cancer Society website	5 websites (QuitNet, Smoke Clinic, V-CC, ProChange, Oregon Center for Applied Sciences) providing tailored, interactive service including peer support, role models, emails, expert advice, information provision and chat room	3 months	RCT-2 groups: (i) Access to static Internet site containing downloadable self-help booklets; (ii) Access to intervention websites free of charge	No difference in cessation rates between (i) and (ii) but in (ii) greater quit rates for those assigned to highly utilised websites

Reference	Sample and setting	Recruitment	Intervention	Follow-up	Design	Main Findings
Prochaska (2008)	136 smokers employed at university; USA; Not required to intend to stop smoking	Mail, email, phone calls	ProChange LifeStyle: Multi-behaviour targeted online intervention programme involving assessment and tailored feedback on basis of stage of change model; fully automated	6 months	RCT-3 groups: (i) brief health risk intervention (based on stages of change);(ii) as (i) + three face-to-face motivational interviewing sessions (iii) as (i) + ProChange intervention	No difference between groups
Strecher (2005)	3971 smokers from general population UK and ROI; Required to intend to stop smoking	Internet website provided in support material of NRT purchased over the counter	CQ Plan: Initial assessment, tailored newsletters, support messages sent by email (based on cognitive-behavioural methods); identification of friend who received tailored email advice to support quitter; fully automated	6, 12 weeks	RCT-2 groups: (i) Web- based control condition that provided similar information but was not tailored (ii) CQ Plan intervention	Significantly greater abstinence rates in (ii) than (i) at follow- up
Swartz (2006)	351 smokers recruited through worksites; USA; Required to intend to stop smoking	Posters, brochures, work intranet website, email and newsletter	1-2-3 Smokefree: Demographically personalised programme providing information, videos to simulate social modelling, tailored quit plan and motivational tips; fully automated	90 days	RCT-2 groups: (i) Waitlist control (ii) 1-2-3 Smokefree intervention	Significantly greater abstinence rates in (ii) than (i) at follow- up

		Popu	lation D	emographics	Λ		Treatment	
Reference	Age (yrs)	Female (%)	White (%)	High School Education ⁺ (%)	Smoking (cig/day)	Length	Intervention Complexity [#]	Control Complexity*
An (2008)	20	72.9	-	100	1.9	20 weeks	high	low
Brendryen (2008a)	36.2	50.3	-	-	18.2	1 year	high	low
Brendryen (2008b)	39.6	50	-	-	17.1	1 year	high	low
Japuntich (2006)	40.8	54.9	79.1	96.8	21.6	90 days	high	medium
McKay (2008)	~42	70.5	86.6	93.1	~21	Not provided	high	high
Munoz (2006)	36.7	54.4	69	Unclear	21.6	8 weeks	high	high
Patten (2006)	15.7	49.6	88	82	9.9	6 months	high	high
Pike (2007)	41	70	-	-	21	Not provided	high	low
Prochaska [†] (2008)	-	-	-	-	-	6 months	high	high
Strecher (2005)	36.9	56.5	-	-	23.5	10 weeks	high	medium
Swartz (2006)	~42	51.9	82.1	-	~19	Not provided	high	-

Table 3: Study samples and treatment characteristics

⁺Schooling until at least 14 years of age; [#]Refers to number of materials used in the intervention, the more complex, the more materials used (e.g. chat rooms, tailored feedback, quizzes etc); *Level of sophistication of treatment in control relative to intervention group (N.B. in cases of more than two study groups, control group refers to group most closely matching intervention group); [^]Where necessary computed indirectly from provided data; [†] Data come from sub-group in study for which no separate characteristics were provided

Figure Legend

Figure 1: Forest plot; RR - Rate ratio; 95% CI - 95% confidence interval

Figure 2: Forest plot; RR - Rate ratio; 95% CI - 95% confidence interval

Figure 3: Forest plot; RR - Rate ratio; 95% CI - 95% confidence interval

Figure 4: Forest plot; RR - Rate ratio; 95% CI - 95% confidence interval; [#]Study results in Munoz et al (2006) are presented separately for the intervention of an English and Spanish websites but for the purpose of this review collapsed across studies

Figure 1: Comparison with minimal, non-interactive, untailored condition

Study or sub-category	Treatment n/N	Control n/N	RR (random) 95% Cl	RR (random) 95% Cl	Year
04 Molting list costrol					
Swertz et el	21/171	9/190		2 46 11 16 5 211	2006
Subtotal (95% CI)	21/1/1	190		2.40 [1.10, 3.21]	2006
Total events: 21 (Treatment)	171 9 (Control)	180		2.46 [1.16, 5.21]	
Test for beterogeneity: not s	n, s (control) applicable				
Test for overall effect: $Z = 2$	2.34 (P = 0.02)				
02 Printed booklet/email cont	rol condition				
An et al	104/257	60/260	_ _ _	1.75 [1.34, 2.29]	2008
Brendryen et al 1	74/197	48/199		1.56 [1.15, 2.11]	2008s
Brendryen et al 2	29/144	10/146		2.94 [1.49, 5.81]	2008b
Subtotal (95% CI)	598	605		1.77 [1.39, 2.27]	
Total events: 207 (Treatmen	t), 118 (Control)				
Test for heterogeneity: Chi2	= 2.82, df = 2 (P = 0.24), l ² = 29	9.1%			
Test for overall effect: Z = 4	.59 (P < 0.00001)				
03 Web-based control cond	ition				
Strecher et al	400/1991	315/1980	-	1.26 [1.10, 1.44]	2005
Pike et al	594/5404	114/1047	÷.	1.01 [0.84, 1.22]	2007
Subtotal (95% Cl)	7395	3027	+	1.14 [0.92, 1.42]	
Total events: 994 (Treatmen	t), 429 (Control)				
Test for heterogeneity: Chi2	= 3.59, df = 1 (P = 0.06), l² = 72	2.1%			
Test for overall effect: Z = 1	.18 (P = 0.24)				
03a Web-based control cor	dition (high usage websites)				
Strecher et al	400/1991	315/1980	-	1.26 [1.10, 1.44]	2005
Pike et al	264/2158	114/1047		1.12 [0.91, 1.38]	2007
Subtotal (95% Cl)	4149	3027	•	1.22 [1.09, 1.37]	
Total events: 664 (Treatmen	t), 429 (Control)				
Test for heterogeneity: Chi2	= 0.87, df = 1 (P = 0.35), l ² = 0°	%			
Test for overall effect: Z = 3	.47 (P = 0.0005)				
		0.1	0.2 0.5 1 2 5	10	

0.1 0.2 0.5 1 2

Favours control Favours treatment

Figure 2: Comparison with minimal, non-interactive, untailored condition-6 months abstinence

Study or sub-category	Treatment n/N	Control n/N	RR (random) 95% Cl	RR (random) 95% Cl	Year			
An et al	104/257	60/260	-	1.75 [1.34, 2.29]	2008			
Brendryen et al 1	73/197	43/199	−	1.71 [1.24, 2.36]	2008a			
Brendryen et al 2	34/144	10/146	_-	3.45 [1.77, 6.71]	2008b			
Total (95% Cl)	598	605	•	1.93 [1.44, 2.60]				
Total events: 211 (Treatmen	it), 113 (Control)		-					
Test for heterogeneity: Chi2	= 3.77, df = 2 (P = 0.15), I ² = 47	.0%						
Test for overall effect: Z = 4	4.35 (P < 0.0001)							
			0.1 0.2 0.5 1 2 5	10				
		Favours control Favours treatment						

Figure 3⁻ Comparison with behavioural treatment

Study or sub-category	Treatment n/N	Control n/N	RR (random) 95% Cl	RR (random) 95% Cl	Year
01 Interactive internet interve	ention vs. face-to-face counsel	ling			
Patten et al	4/70	8/69	_	0.49 [0.16, 1.56]	2006
Prochaska et al	8/36	10/29		0.64 [0.29, 1.42]	2008
Subtotal (95% Cl)	106	98		0.59 [0.31, 1.14]	
Total events: 12 (Treatment) Test for heterogeneity: Chi ² Test for overall effect: Z = 1	, 18 (Control) = 0.14, df = 1 (P = 0.70), I² = 0% .58 (P = 0.11)	5			
02 Interactive internet interve	ention+standard behavioural tre	atment vs.standard behaviou	ral treatment alone		
Japuntich et al	21/140	17/144	_ 	1.27 [0.70, 2.31]	2006
Prochaska et al	8/36	6/36		1.33 [0.51, 3.46]	2008
Subtotal (95% Cl)	176	180		1.29 [0.78, 2.13]	
			0.2 0.5 1 2 5	10	
			Favours control Favours treatme	nt	
Fiaure 4: Compa	arison with enhar	nced interactive	Internet intervention [#]		
Study	Treatment	Control	RR (random)	RR (random)	
or sub-category	n/N	n/N	95% CI	95% CI	Year
Munoz et al	41/281	57/287		0.73 [0.51, 1.06]	2006
McKay et al	45/1159	44/1159	-+-	1.02 [0.68, 1.54]	2008
Total (95% CI) Total events: 86 (Treatment) Test for heterogeneity: Chi ² : Test for overall effect: Z = 0	1440 , 101 (Control) = 1.41, df = 1 (P = 0.24), I ² = 28 .94 (P = 0.35)	1446 9%	•	0.86 [0.62, 1.18]	

0.1 0.2 0.5 1 ż 5 10

Favours control Favours treatment