CADTH RAPID RESPONSE REPORT: SUMMARY WITH CRITICAL APPRAISAL

e-Therapy Interventions for the Treatments of Substance Use Disorders and Other Addictions: A Review of Clinical Effectiveness

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#### Authors: Chuong Ho, Melissa Severn

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#### **Context and Policy Issues**

Alcohol, cannabis, and other substances, [such as stimulants (cocaine, amphetamines), opioids, and sedatives] are among the substances that are the most commonly misused in Canada.<sup>1</sup> This misuse can lead to short- and long-term dependence, disability, mental health issues, and social problems.<sup>1</sup> At least 20% of alcohol is consumed at a level considered to be high-risk drinking.<sup>2,3</sup> The prevalence of cannabis use among the general Canadian population was 12.3% in 2015, and approximately 10% of those who used cannabis developed a cannabis use disorder.<sup>4,5</sup> While not necessarily representative of misuse, in 2013, approximately 22.0% of Ontario adults participated in gambling on a weekly basis, with lottery purchase being the most common.<sup>6-8</sup> It is estimated that 2.5% of adults in Ontario participate in gambling at a level that is moderately or severely problematic.<sup>9</sup>

Despite the positive impact that mental health and addiction services can have in reducing problematic substance use and addiction problems, overcrowded programmes, time restraints (i.e. time out of ones day to travel to and participate in treatment), financial barriers, and fear of stigma, are a few of the barriers to accessing traditional face-to-face care, particularly in rural areas. Interventions for the treatment of problematic substance use and gambling usually include psychosocial interventions such as cognitive behavioral therapy (CBT), motivational interviewing (MI), behavioral self-management (BSM), and personalized normative feedback (PNF).<sup>10</sup> E-therapy, using information technology such as internet- and mobile-based interventions to deliver these psychosocial interventions, has been introduced as a treatment option with the hope of improving access to treatment, reducing time constraints, and reducing costs.<sup>11,12</sup> In general, these treatment modalities include structured self-help using online written materials, and/or audio/video files for the participant to use without assistance (pure self-help), or with assistance from therapists by phone, video or emails (therapist-guided). Therapist guidance can be synchronous (the therapist and the patient are speaking or interacting in real time), or asynchronous (the communication is not in real time).

As e-therapies have the potential to increase access to care, it is important to understand their effectiveness. This Rapid Response report aims to review the comparative effectiveness of therapist-guided e-therapy interventions versus other options for the treatment of adults with substance use disorders and other addictions.

#### **Research Question**

What is the clinical effectiveness of e-therapy for the treatment of patients with substance use disorders and other addictions?

#### **Key Findings**

Evidence on e-therapy for the treatment of substance use disorders and other addictions was from a small number of systematic reviews (SRs) and randomized controlled trials (RCTs). Findings consistently showed that therapist-guided e-therapy was superior to no treatment and wait list in reducing alcohol consumption or cannabis use, and the effect was

small. Therapist-guided e-therapy was found to be equivalent to no treatment and wait list for patients with gambling addiction. With respect to substances, evidence was limited to the treatment of problematic alcohol and cannabis use and it is therefore unclear if the results generalize to the misuse of other substances. The evidence on gambling was limited to those who participated in online video poker.

#### **Methods**

A limited literature search was conducted on key resources including Medline via OVID, PsycINFO via OVID, PubMed, The Cochrane Library, University of York Centre for Reviews and Dissemination (CRD) databases, Canadian and major international health technology agencies, as well as a focused Internet search. Methodological filters were applied to limit retrieval to health technology assessments, systematic reviews, meta-analyses, and randomized controlled trials. The search was also limited to English language documents published between Jan 1, 2013 and May 23, 2018.

Rapid Response reports are organized so that the evidence for each research question is presented separately.

#### Selection Criteria and Methods

One reviewer screened citations and selected studies. In the first level of screening, titles and abstracts were reviewed and potentially relevant articles were retrieved and assessed for inclusion. The final selection of full-text articles was based on the inclusion criteria presented in Table 1.

Population	Adult patients with diagnosed substance use disorders or with a diagnosed addiction disorder (alcohol, drugs, and gambling) with or without co-morbid mental health conditions. Subgroups of interest: military, paramilitary, veterans		
Intervention	Therapist-guided e-therapy: e-therapy interventions (that are not solely video chat or telephone-based) that include therapist contact such as online/internet/web and mobile based therapy		
Comparator	In person treatment, video chat or videoconference therapy, telehealth, wait list, treatment as usual		
Outcomes	Improvements or reductions in symptoms, improved functioning, drop-out rates/attrition rates, synchronicity of therapist guidance		
Study Designs	Heath technology assessments (HTAs), systematic reviews (SRs), meta-analyses (MAs), randomized controlled trials (RCTs)		

#### **Table 1: Selection Criteria**

#### **Exclusion Criteria**

Articles were excluded if they did not meet the selection criteria outlined in Table 1, they were duplicate publications, or were published prior to 2013. Studies that examined etherapy in general without separating therapist-guided and pure self-help strategies, or studies that examined pure self-help, were excluded. Trials that were included in a reported SR were excluded. Smoking cessation was not included in this review.



#### Critical Appraisal of Individual Studies

The included systematic reviews (SR) and clinical trials were critically appraised using the AMSTAR II,<sup>13</sup> and Downs and Black<sup>14</sup> instruments, respectively. Summary scores were not calculated for the included studies; rather, a review of the strengths and limitations of each included study were described.

#### **Summary of Evidence**

#### Quantity of Research Available

A total of 499 citations were identified in the literature search. Following screening of titles and abstracts, 444 citations were excluded and 55 potentially relevant reports from the electronic search were retrieved for full-text review. No potentially relevant publication was retrieved from the grey literature search. Of these potentially relevant articles, 51 publications were excluded for various reasons, while four publications met the inclusion criteria and were included in this report. Appendix 1presents the PRISMA flowchart of the study selection.

#### Summary of Study Characteristics

A detailed summary of the included studies is provided in Appendix 2.

#### Study Design

Two SRs with meta-analysis<sup>15,16</sup> and two clinical trials<sup>17,18</sup> were included in the review. One SR performed literature search until 2013 and included four relevant RCTs (total 16 studies),<sup>15</sup> one SR performed literature search until 2012 and included four relevant RCTs (total 10 studies); the range of dates of the literature searches were not reported.<sup>16</sup> Both included clinical trials are single center, unblinded randomized controlled trials (RCTs).

#### Country of Origin

One SR was performed in the Netherlands, Canada, UK and Germany,<sup>15</sup> and one in Australia, the Netherlands, and Germany.<sup>16</sup> One RCT was performed in France<sup>17</sup> and one in Switzerland.<sup>18</sup>

#### Patient Population

One SR included 5,612 adults with high-risk alcohol use,<sup>15</sup> and one SR included 4,125 participants of all ages with cannabis addiction or abuse.<sup>16</sup> One clinical trial included 1,122 adults with online gambling problems who did not seek treatment,<sup>17</sup> and one clinical trial included 308 adults with cannabis addiction.<sup>18</sup> No studies involved people with comorbid mental health conditions. None of the included studies examined a military, paramilitary, or veteran population.

#### Interventions and Comparators

One SR compared e-therapy (web-based interventions based on CBT, MI, trans-theoretical model of change, and PNF) to no treatment (assessment only, wait list, or alcohol information brochure).<sup>15</sup> One SR compared e-therapy (computer and internet-based interventions based on CBT and MI) to no treatment (assessment only, wait list).<sup>16</sup> One RCT compared internet-delivered CBT with therapist guidance to wait list (synchronicity not

specified),<sup>17</sup> and one RCT compared web-based interventions based on CBT, MI, and BSM (therapist guidance was synchronous).<sup>18</sup>

#### Outcomes

The difference in alcohol consumption after treatment between the two groups was reported on in one SR<sup>15</sup> and the difference in cannabis use between the two groups in another SR.<sup>16</sup> One clinical trial reported the difference from baseline in gambling severity using Problem Gambling Index Score (PGSI – a reduction in score means lower levels of problem gambling) and attrition rate after 6 weeks of treatment,<sup>17</sup> and one clinical trial reported difference from baseline in frequency and quantity of cannabis use and attrition rate after 3 months of treatment.<sup>18</sup>

#### Summary of Critical Appraisal

Details of the strengths and limitations of the included studies are summarized in Appendix 3.

The included SRs<sup>15,16</sup> provided an a priori design and performed a systematic literature search; procedures for the independent duplicate selection and data extraction of studies were in place, a list of included studies and characteristics were provided, a list of excluded studies was not provided. Quality assessment of the included studies was performed and used in formulating conclusions, and publication bias was assessed in both SRs. Both SRs were based mainly on clinical trials which were heterogeneous with respect to the interventions examined (e.g., components and length of e-therapy, amount and type of therapist's support), which may affect the accuracy and reliability of the findings.

The included clinical trials<sup>17,18</sup> were RCTs, the hypotheses were clearly described, the method of selection from the source population and representation were described, losses to follow-up were reported, main outcomes, interventions, patient characteristics, and main findings were clearly described, and estimates of random variability and actual probability values were provided. Both trials performed calculations to determine that the trial was powered to detect a clinically important effect. Patients and assessors were not blinded to treatment assignment in both trials which may have impacted the objectivity of the outcomes assessments. Overall, the included studies had good internal validity; their external validity was limited to people with alcohol, cannabis, and non-treatment seeking poker gambling addiction.

#### Summary of Findings

Details of the findings of the included studies are provided in Appendix 4.

### What is the clinical effectiveness of e-therapy for the treatment of patients with substance use disorders and other addictions?

One SR examined the efficacy of internet-delivered interventions for alcohol addiction.<sup>15</sup> The difference in alcohol consumption between the e-therapy with therapist guidance and no treatment groups at the post-treatment follow-up was statistically significant (with the e-therapy group having less consumption) and the effect of guided e-therapy was found to be small. The authors concluded that internet-delivered interventions were effective in reducing alcohol consumption and that the effect was small.

One SR<sup>16</sup> and one RCT<sup>18</sup> examined the efficacy of e-therapy for cannabis addiction. The SR<sup>16</sup> found that the difference in cannabis consumption between the e-therapy with therapist guidance and no treatment groups at the post-treatment follow-up was statistically significant (with the e-therapy group having lower consumption) and the effect of guided e-therapy was found small. The authors concluded that internet interventions appeared to be effective in reducing cannabis use. The RCT found that guided e-therapy reduced cannabis use on average by one day per week, and the difference was statistically significant, and 23.7% of participants in e-therapy with guidance completed treatment.<sup>18</sup> The authors concluded that web-based interventions were an effective alternative to no treatment or wait list.

One RCT examined the efficacy of e-therapy for the treatment of gambling addiction.<sup>17</sup> The reduction in PGSI score at post-treatment versus baseline was similar in the guided e-therapy group and the no treatment group. The reduction from baseline was larger in the pure self-help e-therapy than in the group with guidance. However, a limited number of participants completed the post-treatment PGSI; the drop-out rates were 97.3 in the group with guidance and 83% in wait list group. The authors concluded that there was lack of significant difference in efficacy between e-therapy and no treatment, and that guidance could have adversely affected those who had not sought help.

#### Limitations

Evidence on therapist-guided e-therapy for the treatment of adults with substance use disorders was based on a small number of SRs and RCTs. The accuracy of estimates from SRs was affected by the heterogeneity in the e-therapy treatments, lack of details on the components of e-therapy strategies, and undetermined amount and type of therapist support (e.g., telephone, email). Together with the lack of details on synchronicity of therapist contact in the included SRs, this is a major limitation that affects the precision of the findings. In most studies, the use of a waitlist as a comparator instead of an active treatment comparator might have led to an overestimate of the treatment effect of e-therapy. Since the included studies either did not include or did not perform subgroup analyses based on a military, paramilitary, or veteran population, the generalizability to this population is unclear. Additionally, it is unclear whether the results of the studies generalize to a population using substances other than cannabis or alcohol, as they were the only substances examined in the included studies.

#### **Conclusions and Implications for Decision or Policy Making**

Limited evidence from the included studies showed that therapist-guided e-therapy reduced problematic alcohol consumption or cannabis use, and that the effect was small. Therapist-guided e-therapy was found to be equivalent to no treatment and wait list for patients with gambling addiction, however the drop-out rate was high, and authors concluded that guidance could have adversely affected those who had not sought help. The accuracy of the evidence on the clinical effectiveness of guided e-therapy was affected by the heterogeneity in the e-therapy treatments and lack of consistency in amount and type of therapist support. A narrative review of SRs on problematic alcohol use agreed that more research on the impact of different levels of guidance is needed to clarify its optimal effect.<sup>19</sup> In agreement with our findings, Internet interventions in general (without guidance status specification) were also found to have a small but significant effect compared to no treatment in reducing alcohol, opioid, cocaine, amphetamine, and methamphetamine use in various SRs.<sup>20,21</sup>

While this review did not do an in-depth search or analysis of cost-effectiveness studies, some cost information was identified, and from a healthcare provider perspective, e-therapy plus treatment-as-usual (TAU) with a counsellor's involvement was found to be likely to be more cost-effective than TAU alone.<sup>22</sup> With the threshold for cost-effectiveness generally based on the healthcare budget available, the generalization of these findings to a Canadian context is limited.

The majority of studies on e-therapy for the treatment of substance use disorders examined the efficacy of the intervention in general, they did not separate therapist-guided from pure self-help, and were therefore not included in our review. While it is likely that therapist-guided e-therapy interventions are effective, larger comparative RCTs with consistent reporting would further confirm the clinical effectiveness of therapist guided e-therapies for the treatment of substance use disorders and other addictions.

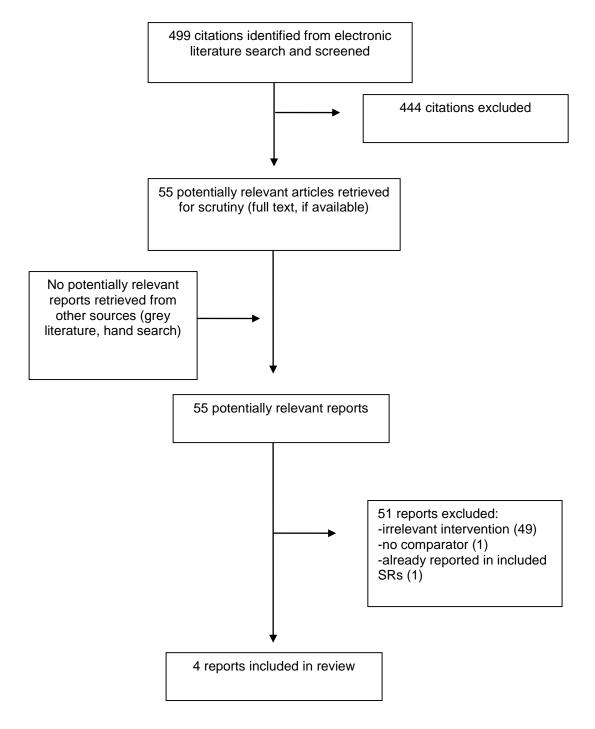
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### **Appendix 1: Selection of Included Studies**





### **Appendix 2: Characteristics of Included Publications**

### Table 2: Characteristics of Included Systematic Reviews

First author, Year, Country	Objectives Intervention Comparators Literature Search Strategy	Inclusion Criteria	Exclusion Criteria	Number of Studies Outcomes
Riper, <sup>15</sup> 2014, Germany, Netherlands, Canada, Australia	<ul> <li>Objectives: "Internet interventions for curbing adult alcohol misuse have been shown effective. Few meta-analyses have been carried out, however, and they have involved small numbers of studies, lacked indicators of drinking within low risk guidelines, and examined the effectiveness of unguided self-help only. We therefore conducted a more thorough meta- analysis that included both guided and unguided interventions" (p 1)</li> <li>Intervention: web-based interventions based on CBT, MI, trans-theoretical model of change, and PNF.</li> <li>Comparators: no treatment (assessment only, wait list or alcohol information brochure)</li> <li>Literature search strategy: "We conducted literature searches up to September 2013 in the following bibliographic databases: MEDLINE, PsycINFO, Science Citation Index Expanded, Social Sciences Citation Index, Arts and Humanities Citation Index, CINAHL, PUBMED and EMBASE, using key words and text words" (p 2)</li> </ul>	"Randomised controlled trials were included that (1) compared a web-based intervention with a control group (in an assessment only, waitlisted or alcohol information brochure control condition); (2) included a low- intensity self-help intervention that the participant could perform on a computer or mobile phone, with or without guidance from a professional; (3) assessed alcohol drinking behaviour in terms of quantity consumed as a primary outcome measure; (4) studied adults aged 18 or older; (5) included alcohol drinkers who exceeded local guidelines for low-risk drinking" (p 2)	Studies not fulfilling exclusion criteria	16 RCTs (4 RCTs with therapist guidance) Efficacy: Effect size of intervention on level of alcohol consumption at post treatment, using AUDIT or FAST score - Therapist- guided - Pure self-help (findings not reported in this review)
Tait, <sup>16</sup> 2013, Australia, Netherlands, Germany	<b>Objectives:</b> "the aim of this review and meta- analysis was to assemble evidence on the effectiveness of computer	"Studies were included in the metaanalysis if they (1) applied a randomized controlled design, (2) tested the effect of an Internet or	Studies not fulfilling exclusion criteria	10 RCTs (4 RCTs with therapist guidance) Efficacy: Effect size of

First author, Year, Country	Objectives Intervention Comparators Literature Search Strategy	Inclusion Criteria	Exclusion Criteria	Number of Studies Outcomes
	and Internet-based interventions in decreasing the frequency of cannabis use and to provide an estimate of the magnitude of that effect" (p 297) Intervention: computer and internet-based interventions based on CBT, MI Comparators: no treatment (assessment only, wait list) Literature search strategy: "In September, 2012, we searched Medline, PubMed, PsychINFO (1806–2012) and Embase (1980–2012). The search terms were (substance related disorders or addiction, or abuse, or dependence or illicit) and (cannabis or marijuana or marihuana or hashish) and (Internet or web or online or computer or CD ROM) and (prevention or treatment or intervention)" (p 297)	computer-delivered intervention (either with or without additional therapeutic guidance) aiming at prevention, indicated prevention or treatment of substance use, (3) reported cannabis use as (one of) the outcome measure(s) and (4) provided usable data to perform the meta-analysis" (p 297)		intervention on level of cannabis use at post treatment, using different measures of cannabis use. - Therapist- guided - Pure self-help (findings not reported in this review)

AUDIT = Alcohol Use Identification Test; CBT = cognitive behavioral therapy; FAST = Fast Alcohol Screening Test; MI = motivational interviewing; PNF = personalized normative feedback; RCT = randomized controlled trial

First Author, Year, Country	Study Design Objectives	Intervention Comparators	Patients	Main Study Outcomes
Luquiens, <sup>17</sup> 2016, France	RCT "The aim of this study was to assess the efficacy of three modalities of Internet-based psychotherapies with or without guidance, compared to a control condition, among problem gamblers who play online poker" (p 1)	Computer- delivered CBT program emailed weekly by a trained psychologist with personalized guidance No treatment (wait list)	1122 adults with PGSI score ≥5 who did not seek treatment -301 with e-therapy with guidance -264 wait list -557 email or CBT book without guidance (findings not reported in this review)	Efficacy: Difference from baseline in PGSI score at 6 weeks Attrition rates
Schaub, <sup>18</sup> 2015, Switzerland	RCT "To test the efficacy of a Web-based self-help intervention with and without chat counseling—Can Reduce—in reducing the cannabis use of problematic cannabis users as an alternative to outpatient treatment services" (p 1)	Web-based self- help intervention, in combination with or without tailored chat counseling, based on CBT, MI, and BSM No treatment (wait list)	308 adults -114 e-therapy with guidance -93 wait list -101 e-therapy without guidance (findings not reported in this review)	Efficacy: Difference from baseline in frequency and quantity of cannabis use at 3 months Attrition rates

#### **Table 3: Characteristics of Included Clinical Studies**

BSM = behavioral self-management; CBT = cognitive behaviour therapy; MI = motivational interviewing; PGSI = Problem Gambling Severity Index; RCT = randomized controlled trial



### **Appendix 3: Critical Appraisal of Included Publications**

### Table 4: Strengths and Limitations of Systematic Reviews and Meta-Analyses using AMSTAR II<sup>13</sup>

Strengths	Limitations			
Riper <sup>15</sup>				
<ul> <li>a priori design provided</li> <li>independent studies selection and data extraction procedure in place</li> <li>comprehensive literature search performed</li> <li>list of included studies, studies characteristics provided</li> <li>quality assessment of included studies provided and used in formulating conclusions</li> <li>assessment of publication bias performed</li> <li>conflict of interest stated</li> </ul>	<ul> <li>list of excluded studies not provided</li> <li>heterogeneity across trials in e-therapy programs (content, lengths, amount and types of therapist support)</li> </ul>			
Tait <sup>16</sup>				
<ul> <li>a priori design provided</li> <li>independent studies selection and data extraction procedure in place</li> <li>comprehensive literature search performed</li> <li>list of included studies, studies characteristics provided</li> <li>quality assessment of included studies provided and used in formulating conclusions</li> <li>assessment of publication bias performed</li> <li>conflict of interest stated</li> </ul>	<ul> <li>list of excluded studies not provided</li> <li>heterogeneity across trials in e-therapy programs (content, lengths, amount and types of therapist support)</li> </ul>			

### Table 5: Strengths and Limitations of Randomized Controlled Trials using Downs and Black<sup>14</sup>

Strengths	Limitations			
Luquiens <sup>17</sup>				
<ul> <li>randomized controlled trial</li> <li>hypothesis clearly described</li> <li>method of selection from source population and representation described</li> <li>loss to follow-up reported</li> <li>main outcomes, interventions, patient characteristics, and main findings clearly described</li> <li>estimates of random variability and actual probability values provided</li> <li>power calculation to detect a clinically important effect performed</li> </ul>	<ul> <li>patient and assessor not blinded to patient treatment assignment.</li> </ul>			
Scha	aub <sup>18</sup>			
<ul> <li>randomized controlled trial</li> <li>hypothesis clearly described</li> <li>method of selection from source population and representation described</li> <li>loss to follow-up reported</li> <li>main outcomes, interventions, patient characteristics, and main findings clearly described</li> <li>power calculation to detect a clinically important effect performed</li> <li>estimates of random variability and actual probability values provided</li> </ul>	<ul> <li>patient and assessor not blinded to patient treatment assignment.</li> </ul>			



### Appendix 4: Main Study Findings and Author's Conclusions

#### Table 6: Summary of Findings of Included Studies

Main Study Findings	Author's Conclusion			
Alcoh	ol use			
Riper <sup>15</sup> (systematic review)				
Difference in alcohol consumption between e-therapy and no treatment at post-treatment (effect size g <sup>a</sup> ) Therapist-guided (data from 4 studies) g = 0.23 (small effect); 95% CI 0.05, 0.41 (Adherence rate was not reported for interventions with therapist	"Internet interventions are effective in reducing adult alcohol consumption and inducing alcohol users to adhere to guidelines for low-risk drinking. This effect is small but from a public health point of view this may warrant large scale implementation at low cost of Internet interventions for adult alcohol misuse" (p 1)			
guidance)				
Cannal	bis use			
Tait <sup>16</sup> (syster	matic review)			
Difference in cannabis use between e-therapy and no treatment at post-treatment (effect size g <sup>a</sup> ) Therapist-guided (data from 4 studies) g = 0.17 (small effect); 95% CI 0.07, 0.26	<i>"Internet and computer interventions appear to be effective in reducing cannabis use in the short-term albeit based on data from few studies and across diverse samples" (p 295)</i>			
Schaub <sup>18</sup> (RCT)				
Change in number of cannabis use days per week compared to baseline (mean) Therapist-guided: 1.4; 95% CI 0.07, 0.61; <i>P</i> 0.02 Change in number of cannabis use days per week compared to	"Web-based self-help interventions supplemented by brief chat counseling are an effective alternative to face-to-face treatment" (p 1)			
wait list (mean) Therapist-guided: 1.0; 95% CI -0.07, 0.47; P 0.03				
23.7% of participants in e-therapy with guidance completed treatment				
Gam	bling			
Luquiens	s <sup>17</sup> (RCT)			
Difference in PGSI score at post-treatment compared to baseline (mean; SD) E-therapy with guidance: -1.64 (3.9)	"Guidance could have aversively affected problem gamblers who had not sought help. Despite the lack of significant difference in efficacy between groups, this naturalistic trial provides a basis for the development of future Internet-based			
Wait list: -1.32 (3.1) (in the email group without guidance, the difference from baseline was larger than in the group with guidance; statistical significance not reported)	trials in individuals with gambling disorders" (p 1)			
Dropout rate E-therapy with guidance: 97.3% Wait list: 83% <sup>a</sup> Effect size g = difference between groups in post treatment alcohol consumption sco				

<sup>a</sup>Effect size g = difference between groups in post treatment alcohol consumption score/SD; 95% CI = 95% confidence interval; CBT = cognitive-behavior therapy; PGSI – Problem Gambling Severity Index score; SD = standard deviation