

VU Research Portal

Internet-based treatment of depression

Cuijpers, P.; Riper, H.; Andersson, G.

published in Current Opinion in Psychology 2015

DOI (link to publisher) 10.1016/j.copsyc.2014.12.026

document version Publisher's PDF, also known as Version of record

Link to publication in VU Research Portal

citation for published version (APA) Cuijpers, P., Riper, H., & Andersson, G. (2015). Internet-based treatment of depression. *Current Opinion in* Psychology, 4, 131-135. https://doi.org/10.1016/j.copsyc.2014.12.026

General rights

Copyright and moral rights for the publications made accessible in the public portal are retained by the authors and/or other copyright owners and it is a condition of accessing publications that users recognise and abide by the legal requirements associated with these rights.

- Users may download and print one copy of any publication from the public portal for the purpose of private study or research.
- You may not further distribute the material or use it for any profit-making activity or commercial gain
 You may freely distribute the URL identifying the publication in the public portal ?

Take down policy If you believe that this document breaches copyright please contact us providing details, and we will remove access to the work immediately and investigate your claim.

E-mail address: vuresearchportal.ub@vu.nl



ScienceDirect



Internet-based treatment of depression Pim Cuijpers^{1,2}, Heleen Riper^{1,2,3} and Gerhard Andersson^{4,5}

A considerable number of randomized controlled trials have shown that Internet-based interventions are effective in the treatment of depression. In the past years a large number of studies are currently focusing on how this knowledge can be used and implemented in mental health care, including preventive services, primary care, specialized mental health care, and general medical care. Furthermore, research on Internet interventions can directly incorporate many new technological applications of the Internet and mobile technology, serious gaming, avatars and automatic emotion recognition, and smartwatches. There is no doubt that this field of research will have a large and lasting impact on mental health care.

Addresses

- ¹ Department of Clinical Psychology, VU University Amsterdam, The Netherlands
- ² EMGO Institute for Health and Care Research, The Netherlands ³ University of Southern Denmark, Department of Telepsychiatry, Denmark
- ⁴ Department of Behavioural Sciences and Learning, Linköping University, Linköping, Sweden
- ⁵ Department of Clinical Neuroscience, Karolinska Institutet, Stockholm, Sweden

Corresponding author: Cuijpers, Pim (p.cuijpers@vu.nl)

Current Opinion in Psychology 2015, 4:131-135

This review comes from a themed issue on Depression

Edited by Christopher G Beevers

For a complete overview see the <u>Issue</u> and the <u>Editorial</u> Available online 30th December 2014

http://dx.doi.org/10.1016/j.copsyc.2014.12.026

2352-250X/O 2014 Elsevier Ltd. All rights reserved.

Introduction

Major depressive disorders constitute a major challenge for health care in the upcoming decades [1]. Major depressive disorders (MDD) are currently ranked fourth worldwide in terms of disease burden, and are expected to rank first in disease burden in high-income countries by the year 2030 [2]. The loss of quality of life for patients and their relatives is substantial, and is worse than in chronic medical conditions such as hypertension, diabetes, and arthritis [3]. Furthermore, the economic costs associated with depressive disorders are huge [4], especially because of production losses, but also by direct and indirect medical costs.

Psychological treatments are considered to be one of the most important interventions to contribute to a reduction

of the disease burden of depressive disorders [5^{••}]. Most patients in Western countries prefer psychological treatment above pharmacological treatments [6]. These treatments have been well developed in recent decades, and several evidence-based psychological treatments exist [7]. However, these treatments have not been scaled up to the extent that they may help reduce the disease burden of mental disorders [8[•]]. Less than half of the people experiencing mental disorders receive treatment, and this is much lower in adolescents, older adults, people with lower socio-economic status, and people from ethnic minorities. Furthermore, the most dominant format in which psychotherapies are delivered is via individual face-to-face contact. This format is much more expensive and time-consuming than other formats, such as guided self-help, and Internet-based treatments. At the same time, most research shows that formats other than individual treatments can also be effective [9,10].

In this paper, we will first present evidence showing that Internet-based guided self-help therapies are effective in the treatment of depression, and the series of studies that have been conducted in recent years to examine how the knowledge that these interventions are effective can be implemented in mental health care. Then we will discuss some recent developments that move away from the 'traditional' Internet interventions, toward mobile health, avatars and serious gaming.

Internet-based interventions are effective in the treatment of depression

A considerable number of randomized controlled trials have shown that Internet-based guided self-help interventions are effective in the treatment of depression $[11^{\circ}]$. In a meta-analysis of studies comparing Internetbased guided self-help with control groups an effect size of d = 0.56 (95% CI: 0.41–0.71) was found. In another meta-analysis only studies among patients with a major depressive disorders were included [12]. This resulted in an effect size of g = 0.78 (95% CI: 0.59–0.96) compared with control conditions, which corresponds with a numbers-needed-to-be-treated (NNT) of two.

These effects correspond well with the effect sizes found for face-to-face psychotherapies for depression [13], such as cognitive behavior therapy and interpersonal psychotherapy. For these treatments effect sizes ranging from 0.5 to 0.8 are found when compared with control groups. Direct comparisons between different types of treatments and treatment formats typically show that there are no major differences between types of therapy for depression or types of treatment format [7]. In a meta-analysis of 21 studies directly comparing faceto-face treatments with guided self-help therapies for depressive and anxiety disorders, no significant difference was found (differential effect size d = 0.02) [9]. We also did not find a significant difference at 3, 6 and 12 months follow-up, nor did we find that the risk of dropping out from treatment was higher (or lower) in the guided selfhelp groups, compared with the face-to-face conditions [13]. These studies focused on guided self-help through the Internet but also through more conventional methods, such as self-help books with telephone support. A more recent meta-analysis focused specifically on trials that compared Internet-based guided self-help treatments with face-to-face therapies (so, no other guided self-help therapies were included) [14^{••}]. The thirteen studies included in this meta-analysis were not only aimed at depression, but also at other problems such as social anxiety disorder, panic disorder, tinnitus, male sexual dysfunction, and body dissatisfaction. The overall difference of all studies was again virtually zero (d = -0.01) and not significant.

Guided Internet-based interventions can be considered as a specific type of guided self-help. Guided self-help can be defined as a psychological treatment in which the patient takes home a standardized psychological treatment protocol and works through it more or less independently [15]. This protocol describes the different steps the patient has to take for applying a generally accepted psychological treatment to himself or herself. The protocol can be presented in book form, or through other media, such as a television, video, computer, or the Internet. Contacts with therapists are not aimed at developing a traditional relationship between therapist and patient, and are only aimed at providing support and, if necessary, added explanation for working through the standardized psychological treatment. Contacts with therapists can be provided through personal contact, by telephone, by e-mail, or by any other available means of communication [15].

Applying Internet-based guided self-help in mental health care

Most of the research on Internet-based guided self-help for depression has been conducted in randomized trials among self-referred depressed people from the community who were recruited by means of advertisements, leaflets and the Internet. But the knowledge that these interventions are effective in the treatment of depression could be implemented in routine mental health care in several ways. Several research groups in Europe, the US and Australia are working on this part of the research agenda [16].

One interesting area where this knowledge can be applied is prevention [13]. Prevention interventions typically have a low intensity nature in terms of access and program intensity, which make Internet-interventions very well suited for this purpose. Several recent and ongoing trials are examining the possibilities of Internet-based interventions to prevent depression [17–20].

Other trials are aimed at examining if and how Internet interventions aimed at the treatment of depression can be integrated in routine primary care [21-23]. Internet-interventions can also be useful in specialized mental health care services, for example during the period between registration and the first contact with a therapist [24], in a 'blended' format where parts of the treatment are delivered in person and other parts through the Internet [25], or as relapse prevention and maintenance treatments [26,27].

A considerable number of trials has also focused on examining whether Internet interventions can be useful for depressed patients who also have a general medical disorder, such as cancer [28,29], diabetes [30,31], and heart disease [32,33]. The application of Internet-based interventions is also examined in areas outside health care, such as school [34], and in occupational settings [35,36].

Innovative applications in e-mental health: self-guided Internet interventions

In the previous paragraph, we saw that Internet-based guided self-help is effective in the treatment of depression that these effects are comparable to those of face-toface treatments, and that many studies are now examining how this knowledge can be applied in mental and public health care. There are, however, also several new and innovative developments in the field of Internet-based interventions that currently have considerable interests among researchers.

One issue is the development of Internet interventions without any kind of human support. As indicated, Internet-based interventions with some kind of professional support have been found to be effective. When there is no support, however, and the patient has to work through the intervention all by himself or herself, the effects are significantly smaller than those of guided interventions [10,37]. In a meta-analysis of unguided Internet-based interventions for depression we found a small effect size of d = 0.28 (95% CI: 0.14–0.42) [38°]. If it would be possible to develop unguided interventions that are as effective as supported interventions, that would be much more efficient and less costly, allowing much broader implementation, because no infrastructure for delivery of the interventions, training and supervision of therapists would be needed. So, several trials continue to examine the effects of self-guided Internet interventions [39–41].

One could also argue that unguided Internet-interventions could be useful in low income and middle income countries without an infrastructure for mental health care. These interventions may be somewhat less effective than guided self-help interventions or face-to-face treatments, they can still contribute considerably to a reduction of the disease burden. If the alternative is that depressed patients receive no treatment whatsoever, then these interventions can reduce the disease burden of depression, especially when the number of people using them is large [42[•]].

Technological innovations and Internet interventions

Technological innovations are an important driving force to new developments in e-health and mobile-health. It is beyond the scope of this paper to give a comprehensive overview of all new developments related to technological innovations, but there are some important examples we should mention.

One important innovation is mobile Internet and smartphones. Real-time assessments [43,44[•]], which are also called 'ecological momentary assessment' or 'experience sampling' offer new possibilities to measure and predict elements of mental health in the natural environment of patients. These types of measurements are often related to various other events [45]. Smartphones also have several sensors, such as the accelerometer that can be used to measure movement or GPS sensors that measure location [46]. The smartphone offers a wealth of (often unknown) indirect data as well. For example with innovative apps the actual use of the phone in terms of phoning, text messaging or Internet or social media use, can also be applied to develop personalized models for changes in depression over time and to predict changes in mood. This allows the development of personalized advice and 'ecological momentary interventions' which are based on the personal information of the user.

But there are other more technologically driven developments. A growing number of randomized trials have shown for example that interventions based on virtual reality (VR) have strong effects on depression [47]. This also holds for an even more recent development, that is, augmented reality (AR), that blends seamlessly virtual and real life components for example in the treatment of arachnophobia. Both VR and AR extend the array of exposure based formats between and interceptive and vivo exposure [48]. One advantage of AR over VR that it is cheaper and still more easy to implement in routine practice. 'Serious gaming' has also been shown to be effective in the treatment of depression in adolescents [49^{••}]. Other technological applications that can be expected to change Internet interventions include the development of avatars, smartwatches and things like 'google glass'. Last but not least, an increased number of programs enable secured videoconferencing between professionals and between therapists and their patients for both consulting and therapeutic interventions with promising results for clinical effectiveness regarding depression treatments [50].

Although it is not clear how exactly these technological developments will change currently existing Internet interventions, it is clear that the impact will be large and that these will have a large impact on this field and the field of mental health treatments in general.

Conclusions

Since their first development in the late 1990s Internetbased treatments of common mental disorders, a considerable number of randomized controlled trials have shown that these interventions are effective in the treatment of depression, especially when they are using a guided self-help format. Although Internet-interventions without professional guidance are less effective than guided interventions, they still may have significant effects on reduction of depressive symptoms. Currently a large number of studies are examining how this knowledge can be used and implemented in mental health care, including preventive services, primary care, specialized mental health care, and general medical care. One interesting research question is if and how unguided interventions can be as effective as guided self-help interventions as this would enable large scale upscaling of low cost mental disorder prevention and treatment initiatives. We also showed that Internet interventions can build directly on the many new technological applications of the Internet and mobile technology, through ecological momentary assessment and intervention, serious gaming, avatars and automatic emotion recognition, and smartwatches.

The field of research on Internet interventions is booming and there are many possibilities for all kinds of innovative interventions and applications. There should be no doubt that this research will have a large and lasting impact on the field mental health care, the therapeutic process and patient-therapist relationships and thus it will change this field considerably. Robust evidence based studies should accompany however new technology based depression treatments before they are implemented in routine care.

Conflict of interest

None declared.

References and recommended reading

Papers of particular interest, published within the period of review, have been highlighted as:

- of special interest
- •• of outstanding interest
- Collins PY, Patel V, Joestl SS, March D, Insel TR, Daar AS et al.: Grand challenges in global mental health. Nature 2011, 475:27-30.

- Mathers CD, Loncar D: Projections of global mortality and burden of disease from 2002 to 2030. PLoS Med 2006, 3:e442.
- Wells KB, Stewart A, Hays RD, Burnam MA, Rogers W, Daniels M, Berry S, Greenfield S, Ware J: The functioning and well-being of depressed patients. Results from the Medical Outcomes Study. JAMA 1989, 262:914-919.
- Smit F, Cuijpers P, Oostenbrink J, Batelaan N, de Graaf R, Beekman A: Excess costs of common mental disorders: population based cohort study. J Ment Health Policy Econ 2006, 9:193-200.
- Emmelkamp PM, David D, Beckers T, Muris P, Cuijpers P, Lutz W
 et al.: Advancing psychotherapy and evidence-based
- et al.: Advancing psychological interventions. Int J Methods Psychiatr Res 2014, 23(Suppl 1):58-91.

This paper gives a nice overview of priorities for research on psychological interventions in general, including Internet-interventions, developed by major experts in the field.

- McHugh RK, Whitton SW, Peckham AD, Welge JA, Otto MW: Patient preference for psychological vs pharmacological treatment of psychiatric disorders: a meta-analytic review. J Clin Psychiatry 2013, 74:595-602.
- Barth J, Munder T, Gerger H, Nuesch E, Trelle S, Znoj H, Juni P, Cuijpers P: Comparative efficacy of seven psychotherapeutic interventions for depressed patients: a network metaanalysis. *PLoS Med* 2013, 10:e1001454.
- Kazdin AE, Blasé SL: Rebooting psychotherapy research and practice to reduce the burden of mental illness. *Perspect Psychol Sci* 2011, 6:21-37.

Excellent analysis of the state-of-the-art in research on psychological treatments in general, focusing on the major strengths and weaknesses of the field, and included an overview of research priorities.

- Cuijpers P, Donker T, van Straten A, Li J, Andersson G: Is guided self-help as effective as face-to-face psychotherapy for depression and anxiety disorders? A systematic review and meta-analysis of comparative outcome studies. *Psychol Med* 2010, 40:1943-1957.
- 10. Andersson G, Cuijpers P: Internet-based and other computerized psychological treatments for adult depression: a meta-analysis. *Cogn Behav Ther* 2009, **38**:196-205.
- Richards D, Richardson T: Computer-based psychological
 treatments for depression: a systematic review and metaanalysis. *Clin Psychol Rev* 2012, 32:329-342.

This is the latest comprehensive review of all randomized trials on computerized therapies for depression.

- 12. Andrews G, Cuijpers P, Craske MG, McEvoy P, Titov N: Computer therapy for the anxiety and depressive disorders is effective, acceptable and practical health care: a meta-analysis and pilot implementation. *PLoS ONE* 2010, **5**:e13196.
- Cuijpers P, Riper H: Internet interventions for depressive disorders: an overview. Span J Psychol 2015. (in press).
- 14. Andersson G, Cuijpers P, Carlbring P, Riper H, Hedman E:
- Internet-based vs. face-to-face cognitive behaviour therapy for psychiatric and somatic disorders: a systematic review and meta-analysis. *World Psychiatry* 2014, **13**:288-295.

All randomized trials directly comparing face-to-face therapies with Internet-based treatments for any disorder are included in this systematic review and meta-analysis. It clearly shows that there is no indication to assume that there are major differences between the two treatment formats.

- Cuijpers P, Schuurmans J: Self-help interventions for anxiety disorders: an overview. Curr Psychiatry Rep 2007, 9:284-290.
- Riper H, Andersson G, Christensen H, Cuijpers P, Lange A, Eysenbach G: Theme issue on E-mental health: a growing field in Internet research. J Med Intern Res 2010, 12:e74.
- Gosling JA, Glozier N, Griffiths K, Ritterband L, Thorndike F, Mackinnon A, Hehir KK, Bennett A, Bennett K, Christensen H: The GoodNight study – online CBT for insomnia for the indicated prevention of depression: study protocol for a randomised controlled trial. *Trials* 2014, 15:56.

- Barrera AZ, Kelman AR, Muñoz RF: Keywords to recruit Spanish- and English-speaking participants: evidence from an online postpartum depression randomized controlled trial. J Med Internet Res 2014, 16:e6.
- Buntrock C, Ebert DD, Lehr D, Cuijpers P, Riper H, Smit F, Berking M: Evaluating the efficacy and cost-effectiveness of web-based indicated prevention of major depression: design of a randomised controlled trial. *BMC Psychiatry* 2014, 14:25.
- Lintvedt OK, Griffiths KM, Sorensen Kostvik AR, Wang CE, Eisemann M, Waterloo K: Evaluating the effectiveness and efficacy of unguided Internet-based self-help intervention for the prevention of depression: a randomized controlled trial. *Clin Psychol Psychother* 2013, 20:10-27.
- Newby JM, Mackenzie A, Williams AD, McIntyre K, Watts S, Wong N, Andrews G: Internet cognitive behavioural therapy for mixed anxiety and depression: a randomized controlled trial and evidence of effectiveness in primary care. *Psychol Med* 2013, 43:2635-2648.
- Høifødt RS, Lillevoll KR, Griffiths KM, Wilsgaard T, Eisemann M, Waterloo K, Kolstrup N: The clinical effectiveness of web-based cognitive behavioral therapy with face-to-face therapist support for depressed primary care patients: randomized controlled trial. J Med Internet Res 2013, 15:e153.
- Mohr DC, Duffecy J, Ho J, Kwasny M, Cai X, Burns MN, Begale M: A randomized controlled trial evaluating a manualized TeleCoaching protocol for improving adherence to a web- based intervention for the treatment of depression. PLOS ONE 2013, 8:e70086.
- 24. Kok RN, van Straten A, Beekman AT, Cuijpers P: Short-term effectiveness of web-based guided self-help for phobic outpatients: randomized controlled trial. *J Med Internet Res* 2014, **16**:e226.
- Kooistra LC, Wiersma JE, Ruwaard J, van Oppen P, Smit F, Lokkerbol J, Cuijpers P, Riper H: Blended vs. face-to-face cognitive behavioural treatment for major depression in specialized mental health care: study protocol of a randomized controlled cost-effectiveness trial. BMC Psychiatry 2014, 14:290.
- Hollandare F, Johnsson S, Randestad M, Tillfors M, Carlbring P, Andersson G, Engstrom I: Randomized trial of Internet-based relapse prevention for partially remitted depression. Acta Psychiatr Scand 2011, 124:285-294.
- 27. Carlbring P, Lindner P, Martell C, Hassman P, Forsberg L, Strom L, Andersson G: The effects on depression of Internetadministered behavioural activation and physical exercise with treatment rationale and relapse prevention: study protocol for a randomised controlled trial. *Trials* 2013, 14:35.
- Mattsson S, Alfonsson S, Carlsson M, Nygren P, Olsson E, Johansson B: U-CARE: Internet-based stepped care with interactive support and cognitive behavioral therapy for reduction of anxiety and depressive symptoms in cancer – a clinical trial protocol. *BMC Cancer* 2013, 13:414.
- 29. Krebber AM, Leemans CR, de Bree R, van Straten A, Smit F, Smit EF, Becker A, Eeckhout GM, Beekman AT, Cuijpers P, Verdonck-de Leeuw IM: Stepped care targeting psychological distress in head and neck and lung cancer patients: a randomized clinical trial. *BMC Cancer* 2012, **12**:173.
- 30. Van Bastelaar K, Pouwer F, Cuijpers P, Riper H, Snoek F: Webbased depression treatment for type 1 and type 2 diabetes patients: a randomized controlled trial. *Diabetes Care* 2011, 34:320-325.
- Nobis S, Lehr D, Ebert DD, Berking M, Heber E, Baumeister H, Becker A, Snoek F, Riper H: Efficacy and cost-effectiveness of a web-based intervention with mobile phone support to treat depressive symptoms in adults with diabetes mellitus type 1 and type 2: design of a randomised controlled trial. BMC Psychiatry 2013, 13:306.
- 32. Habibović M, Cuijpers P, Alings M, van der Voort P, Theuns D, Bouwels L, Herrman JP, Valk S, Pedersen S: Attrition and adherence in a WEB-Based Distress Management Program for Implantable Cardioverter defibrillator Patients

(WEBCARE): randomized controlled trial. *J Med Internet Res* 2014, **16**:e52.

- **33.** Messerli-Bürgy N, Barth J, Berger T: **The InterHerz project a web-based psychological treatment for cardiac patients with depression: study protocol of a randomized controlled trial**. *Trials* 2012, **13**:245.
- Calear AL, Christensen H, Mackinnon A, Griffiths KM: Adherence to the MoodGYM program: outcomes and predictors for an adolescent school-based population. J Affect Disord 2013, 147:338-344.
- 35. Geraedts AS, Kleiboer AM, Wiezer NM, van Mechelen W, Cuijpers P: Short-term effects of a web-based guided self-help intervention for employees with depressive symptoms: randomized controlled trial. J Med Int Res 2014, 16:e121.
- Heber E, Ebert DD, Lehr D, Nobis S, Berking M, Riper H: Efficacy and cost-effectiveness of a web-based and mobile stressmanagement intervention for employees: design of a randomized controlled trial. *BMC Public Health* 2013, 13:655.
- Spek V, Cuijpers P, Nyklíček I, Riper H, Keyzer J, Pop V: Internetbased cognitive behavior therapy for mood and anxiety disorders: a meta-analysis. *Psychol Med* 2007, 37:319-328.
- Cuijpers P, Donker T, Johansson R, Mohr DC, van Straten A,
 Andersson G: Self-guided psychological treatment for depressive symptoms: a meta-analysis. *PLoS ONE* 2011, 6:e21274.

This is a comprehensive meta-analysis of randomized trials comparing self-guided Internet-based interventions for depression with (care-as-usual) control groups. Small, but significant effects are found.

- 39. Gulliver A, Griffiths KM, Christensen H, Mackinnon A, Calear AL, Parsons A, Bennett K, Batterham PJ, Stanimirovic R: Internetbased interventions to promote mental health help-seeking in elite athletes: an exploratory randomized controlled trial. J Med Internet Res 2012, 14:e69.
- Farrer L, Christensen H, Griffiths KM, Mackinnon A: Web-based cognitive behavior therapy for depression with and without telephone tracking in a national helpline: secondary outcomes from a randomized controlled trial. J Med Internet Res 2012, 14:e68.
- Cockayne NL, Glozier N, Naismith SL, Christensen H, Neal B, Hickie IB: Internet-based treatment for older adults with depression and co-morbid cardiovascular disease: protocol

for a randomised, double-blind, placebo controlled trial. *BMC Psychiatry* 2011, **11**:10.

42. Watts SE, Andrews G: Internet access is NOT restricted globally
to high income countries: so why are evidenced based prevention and treatment programs for mental disorders so rare? Asian J Psychiatry 2014, 10:71-74.

Excellent plea to develop Internet-based interventions in low-income countries.

43. Trull TJ, Ebner-Priemer UW: Using experience sampling methods/ecological momentary assessment (ESM/EMA) in clinical assessment and clinical research: introduction to the special section. *Psychol Assess* 2009, **21**:457-462.

44. Wenze SJ, Miller IW: Use of ecological momentary assessment

• in mood disorders research. *Clin Psychol Rev* 2010, **30**:794-804. Excellent overview of research on ecological momentary assessment in depressive disorders.

- 45. Warmerdam L, Riper H, Klein M, van den Ven P, Rocha A, Henriques M, Tousset E, Silva H, Andersson G, Cuijpers P: Innovative ICT solutions to improve treatment outcomes for depression: the ICT4Depression Project. Stud Health Technol Inform 2012, 181:339-343.
- Guiry JJ, van de Ven P, Nelson J, Warmerdam L, Riper H: Activity recognition with smartphone support. Med Eng Phys 2014, 36:670-675.
- Opriş D, Pintea S, García-Palacios A, Botella C, Szamosközi , David D: Virtual reality exposure therapy in anxiety disorders: a quantitative meta-analysis. Depress Anxiety 2012, 29:85-93.
- Baus O, Bouchard S: Moving from virtual reality exposurebased therapy to augmented reality exposure-based therapy: a review. Front Hum Neurosci 2014, 8:112.
- 49. Merry SN, Stasiak K, Shepherd M, Frampton C, Fleming T,
- Lucassen MF: The effectiveness of SPARX, a computerised self help intervention for adolescents seeking help for depression: randomised controlled non-inferiority trial. *BMJ* 2012, 344:e2598.

Fascinating randomized controlled trial showing that a serious game is effective in reducing depressive symptomatology in depressive adolescents.

 Valdagno M1: Telepsychiatry: new perspectives and open issues. CNS Spectr 2014, 2:1-3.