Individually tailored internet-based cognitive behavioural therapy for adolescents, young adults and older adults with anxiety

Kristin Silfvernagel



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- For those who remain, death should never take preced	ence
over life.	

Abstract

Anxiety disorders share the feature of excessive fear, anxiety and related behavioural disturbances. Fear is defined as the emotional response to a real or a perceived imminent threat and anxiety is the anticipation of a future threat. The anxiety disorders covered in this thesis are panic disorder with or without agoraphobia, social phobia, post-traumatic stress disorder, generalized anxiety disorder and anxiety disorder not otherwise specified. Cognitive behavioural treatment protocols are typically designed to target one specific disorder and falls under the definition of disorder-specific cognitive behavioural therapy. It is however unclear if this is the most optimal approach in regards to the high comorbidity between anxiety disorders and depressive disorders. Internet-based cognitive behavioural therapy has in the past generally been disorder-specific and from above mentioned predicament two alternative treatment approaches emerged, the tailored and the transdiagnostic approach that aims to simultaneously treat both principal and comorbid disorders. Previous trials on internet-based cognitive behavioural therapy have targeted adults in general and relatively few target adolescents, young adults and older adults.

The aims of this thesis were to further develop and test the effects of tailored internet-based cognitive behavioural therapy on the basis of age, for adolescents, young adults and older adults. Specifically by developing and testing the effects of individually tailored internet-based cognitive behavioural therapy for adolescents with anxiety and comorbid depressive symptoms and by adapting and testing the effects of individually tailored internet-based cognitive behavioural therapy for young adults and older adults with anxiety and comorbid depressive symptoms. These aims were tested in two pilot effectiveness studies (Paper I and III) and two efficacy randomised controlled trials (Paper II and IV). The results from these four trials showed significant results across all outcome measures with overall moderate to large effect sizes. The tentative conclusion based on these results is that tailoring internet-based cognitive behavioural therapy can be a feasible approach in the treatment of anxiety symptoms and comorbid depressive symptoms for adolescents, young adults and older adults. Despite the positive findings of the studies in this thesis, there is a need for more research examining the acceptability and effectiveness of internet-based cognitive behavioural therapy for adolescents, young adults and older adults with anxiety and depression before implementation on a larger scale.

Keywords: internet-based treatment, cognitive behavioural therapy, anxiety, adolescents, young adults, older adults

Empirical studies

- I. **Silfvernagel, K.**, Gren-Landell, M., Emanuelsson, M., Carlbring, P., & Andersson, G. (2015). Individually tailored internet-based cognitive behavior therapy for adolescents with anxiety disorders: A pilot effectiveness study. *Internet Interventions*, *2*, 297-302.
- II. **Silfvernagel, K.**, Carlbring, P., Kabo, J., Edström, S., Eriksson, J., Månson, L., & Andersson, G. (2012). Individually tailored internet-based treatment for young adults and adults with panic attacks: Randomized controlled trial. *Journal of Medical Internet Research*, *14* (3), e65.
- III. **Silfvernagel, K.**, Wassermann, C., & Andersson, G. (accepted). Individually tailored internet-based cognitive behavioural therapy for young adults with anxiety disorders: A pilot effectiveness study. *Internet Interventions*.
- IV. **Silfvernagel, K.**, Westlinder, A., Andersson, S., Bergman, K., Diaz Hernandez, R., Fallhagen, L., Lundqvist, I., Masri, N., Viberg, L., Forsberg, M-L., Lind, M., Berger, T., Carlbring, P., & Andersson, G. (submitted). Individually tailored internetbased cognitive behaviour therapy for older adults with anxiety and depression: A randomised controlled trial.

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Background

Introduction

Internet-based cognitive behaviour therapy (ICBT) (Andersson, 2009) has emerged as a novel evidence-based treatment for anxiety and mood disorders (Andrews, Cuipers, Craske, McEvoy & Titoy, 2010; Andersson & Cuijpers, 2009). Guided ICBT has been found to be effective for numerous specific disorders such as panic disorder (Carlbring, Westling, Ljungstrand, Ekselius & Andersson, 2001; Klein, Richards & Austin, 2006), generalized anxiety disorder (Robinson et al., 2010; Paxling et al., 2011), social anxiety disorder (Boettcher, Carlbring, Renneberg & Berger, 2013), post-traumatic stress disorder (Lange et al., 2003) and major depression (Andersson & Cuijpers, 2009). One limitation of many previous ICBT trials on anxiety and depression is that they targeted specific disorders and that comorbid disorders could either be affected without being directly addressed or remain undetected and unchanged (Titov, Gibson, Andrews & McEvoy, 2009). Another potential limitation of previous ICBT trials as stated by Carlbring et al. (2006) is that participants with comorbid anxiety and depression disorders are often excluded. Structured or diagnosis-specific ICBT treatments are also limited in that they leave little room for clinician and patient preferences.

One approach to ICBT aims to address these limitations by combining individually tailored treatment according to the participant's needs and symptoms with transdiagnostic components (Carlbring et al., 2011; Johansson et al., 2012; Bergman Nordgren et al., 2014). Previous trials on tailored ICBT have been conducted on an adult population. This thesis examines the effects of tailored ICBT on the basis of age, for adolescents, young adults and older adults.

Anxiety

Anxiety disorders share the feature of excessive fear, anxiety and related behavioural disturbances (American Psychiatric Association, 2013). Fear is defined as the emotional response to a real or a perceived imminent threat and anxiety is the anticipation of a future threat. These states occur simultaneously but vary; fear is associated with autonomic arousal involved in the flight or fight response, thoughts of danger and escape behaviours. Anxiety is more often associated with muscle tension and vigilance in preparation for future danger and avoidant behaviours. Some have a biological vulnerability to react with negative emotions during stress, and to try to alleviate the discomfort the individual feels, they either try to, or escape the situation (Barlow, 2014). Both the biological vulnerability, and the individuals learning history, contribute to the development of anxiety disorders.

The anxiety disorders covered in this thesis are panic disorder with or without agoraphobia, social phobia, post-traumatic stress disorder, generalized anxiety disorder and anxiety disorder not otherwise specified based on the criteria of The Diagnostic and Statistical Manual of Mental Disorders, fourth edition, text revision (DSM-IV-TR; American Psychiatric Association, 2000). Panic attacks are prominently featured within the different anxiety disorders as a fear response and are characterised as a discrete period in which there is a sudden onset of intense apprehension, fearfulness or terror that is often associated with feelings of impending doom. During the attack, symptoms such as accelerated heart rate, sweating, trembling, sensations of shortness of breath, feeling of choking, chest pain, abdominal distress, feeling dizzy, derealisation, fear of losing control, fear of dying, paraesthesia, chills or hot flushes can be present. The different anxiety disorders are distinguished from one another in the types of stimuli that induce fear and anxiety and from the avoidant behaviour present (American Psychiatric Association, 2013). According to the DSM-IV-TR classification system panic disorder with or without agoraphobia is characterised by recurrent unexpected panic attacks from which there is persistent concern (American Psychiatric Association, 2000). Agoraphobia is anxiety about, or avoidance of, places or situations from which escape might be difficult or in which help may not be available in the event of having a panic attack or symptoms of panic. Social phobia is characterised by clinically significant anxiety, provoked by exposure to certain types of social or performance situations, often leading to avoidant behaviour. Posttraumatic stress disorder is characterised by the re-experiencing of an

extremely traumatic event accompanied by symptoms of increased arousal and by avoidance of stimuli associated with the trauma. Generalized anxiety disorder is characterised by at least 6 months of persistent and excessive anxiety and worry. Anxiety disorder not otherwise specified is a category for prominent anxiety or phobic avoidance for individuals that do not meet criteria for any specific anxiety disorder, but the symptoms are severe enough to warrant a diagnosis of an anxiety disorder.

Comorbidity

There is now considerable evidence of a complex and reciprocal relationship between anxiety and depression (Katon & Ciechanowski, 2002; Roy-Byrne et al., 2008). Comorbidity is common in anxiety disorders; between 65-95 percent of individuals who have an anxiety disorder also meet the criteria for another diagnosis (Kovacs & Devlin, 1998). It is most common to have more than one anxiety disorder. Among young people with anxiety disorder 40-60 percent meet the diagnostic criteria for more than one anxiety disorder at the same time (Wenar & Kerig, 2008; Rapee, Schniering & Hudson, 2009). Depression is the second most common comorbid condition of anxiety disorders. Other common comorbid conditions in youths are oppositional defiant disorder, conduct disorder, attention hyperactivity disorder and substance abuse (Rutter et al., 2012; Rapee, Schniering & Hudson, 2009; Costello et al., 2003). Reflecting the bidirectional relationships between psychological and physical illnesses, the prevalence of chronic health conditions increases with age and having a chronic health condition appears to be one of the strongest risk factors for anxiety and depression (Britt et al., 2008). For example, research indicates that the risk of myocardial infarction increases fourfold among adults with symptoms of depression (Hippisley-Cox et al., 1998). Moreover, depressed patients are three times less likely to adhere to recommended medical treatment regimens for their chronic health conditions than non-depressed patients (DiMatteo et al., 2000). Furthermore, while anxiety and depression are major health issues themselves, untreated anxiety and depression are significant issues among older adults because of their association with poor physical health.

Anxiety during adolescence

Epidemiological studies suggest that about 5-19 percent of all children and adolescents have some form of anxiety disorder, which means it is one of the most common psychiatric disorders (James et al., 2015). The median age of onset of any anxiety disorder is 11 years (Kessler et al., 2005; Rickwood & Bradford, 2012). The presentation of anxiety disorders varies with age (James et al., 2015). Young children can present with indistinguishable worries and fears and multiple somatic symptoms (i.e. muscle tension, headache or stomachache) and sometimes behavioral symptoms (i.e. angry outbursts). The latter may be misdiagnosed as oppositional defiant disorder, as the child tries to avoid anxiety-provoking situations. Social phobia normally appears after puberty. Previous research suggests that adolescents with anxiety are a particularly underserved population that often does not receive adequate treatment for their anxiety (Kendall & Peterman, 2015) and many cases of social phobia are first diagnosed more than 20 years after onset (James et al., 2015). Distinguishing between normal, developmentally appropriate worries, fears and shyness from anxiety disorders are one of the diagnostic challenges in this age group. Adolescents typically worry and have fears related to school performance, social competence and health issues. Untreated anxiety disorders have a relatively chronic course and adult studies have found that anxiety disorders often have its onset in childhood or adolescence (Wenar & Kerig, 2008). Adolescents with anxiety disorders also face a range of serious impairments in interpersonal, leisure and academic functioning (Kendall & Peterman, 2015). To operationalize age groups is a challenge, given the heterogeneity of biological, cognitive, emotional, and social maturity in different age groups and due to the fact that there is no consensus at this point in time. Adolescence is however defined by the World Health Organization as between the ages of 10 and 19 years (Patton et al., 2016). In this thesis adolescence is defined as between the ages of 15 and 19, from the age that an individual can give legal consent for trial participation to the age where the individual is no longer considered an adolescent within the child and psychiatric system in Sweden.

Anxiety during young adulthood

Anxiety disorders are among the most common psychiatric disorders in youths (Kendall & Peterman, 2015). Most individuals who develop an anxiety disorder do so by late adolescence or early adulthood (Seligman & Ollendick, 2011). Adolescence is by no means a unified transition from childhood to adulthood (Kendall & Peterman, 2015). Therefore the age group young adult can be considered warranted. In this thesis young adulthood is defined as being aged between 16 and 30, 16 being the youngest age accepted into the youth health care system in Sweden for psychological treatment, and 30 as the upper limit, where most anxiety symptoms have been manifested for the individual. This phase in life is also sometimes described as emerging adulthood; from the late teens to the late twenties when an individual acquires some of the characteristics of adulthood without having reached the milestones that historically define fully fledged adulthood (Patton et al., 2016). Prevalence rates for anxiety in youths vary from 2.4 to 17 percent (Kendall & Suveg, 2006). Anxiety consists of behavioural, somatic, cognitive and emotional features and one of the most prominent behavioural responses to anxiety is avoidance (Kendall & Suveg, 2006). Youth with anxiety normally report an increase in autonomic nervous system activity, perspiration, abdominal pain, a flushed face, gastrointestinal distress and trembling. Many youths with anxiety experience cognitive distress (i.e. rumination and excessive worrying). Social phobia become more prevalent after the adolescent has entered puberty but anxiety disorders with onset in childhood normally continue to persist during adolescence and early adulthood if left untreated (James et al., 2015). During the time period of 1994 to 2006 the proportion of young adults aged 16-29 years who reported severe anxiety, more than doubled in Sweden (Swedish National Board of Health and Welfare, 2013). Anxiety in youth, places the individual at an increased risk for comorbid diagnoses, psychopathology in adulthood and lower adaptive functioning in regards to academic/work performance, relations to peers and family relations (Kendall & Suveg, 2006).

Anxiety in late life

Symptoms of anxiety and depression are common among older adults (Skoog, 2011), but are sometimes mistaken for being natural signs of ageing (Law et al., 2010). However, prevalence studies suggest that about 5-10 percent of adults over the age of 65 fulfil the criteria for an anxiety disorder (Schuurmans & van Balkom, 2011). However, it is argued that the actual prevalence rates may be underestimated in older adults and that many more are likely to experience symptoms of anxiety and depression (O'Connor 2006; Luppa et al., 2012). This is of significant importance since the average life expectancy is set to increase in many countries (Oeppen & Vaupel, 2002; Porensky et al., 2009). The National Board of Health and Welfare in Sweden (2009) reported that 3-5 percent of adults over 65 years met diagnostic criteria for generalized anxiety disorder and that 10-15 percent experienced late life depression. Anxiety and depression are particularly significant issues among older adults because of their association with poor physical health. Research indicates that anxiety and depression in older adults are associated with poorer physical health and the exacerbation of physical illness (Braam et al., 2005; Brenes et al., 2008) as well as reduced quality of life and increased risk of suicide (Grek, 2007). In older adults, the symptoms are not as prominent as in adults in general (Wolitzky-Taylor et al., 2010). For example, older adults reported less physiological symptoms, use other concepts to describe their experiences and derive the symptoms to somatic diseases. Instead of using the concepts included in the criteria for the disorder, such as "concern" or "uncontrollable", older adults rather use the expressions "problem" and "difficulties" which can lead to misunderstandings and, ultimately, missed diagnoses (Bower, Wetherell, Mon & Lenze, 2015). Studies on panic disorder in older adults have shown that older adults react with less panic symptoms and have higher functionality compared with younger adults (Lenze & Wetherell, 2009) and studies on worry content for older adults with or without generalized anxiety disorder show that older adults report a wider variety of worry topics (Diefenbach, Stanley & Beck, 2001). In this thesis older adults is defined as being of age 60 and above.

Cognitive behavioural therapy

In cognitive behavioural therapy (CBT) it is the relationship between thoughts, feelings, behaviours and bodily sensations that are in focus (Wilkinson, 2013). CBT is based on distinctive theoretical frameworks, such as respondent and operant conditioning (Skinner, 1938; Watson & Rayner, 2000) and information-processing cognitive theories (Beck & Clark, 1997; Beck & Haigh, 2014). CBT has its basis in learning psychology, cognitive psychology, and has recently also taken influences from Eastern philosophy; affect theory and dialectics (Westbrook et al., 2011). Modern CBT assumes all these perspectives in the assessment and treatment of mental illness. Based on learning psychology the therapist examines how problem behaviours are perpetuated by contingencies of reinforcement, and based on cognitive psychology, the therapist and patient find how negative thought patterns, presuppositions and rules of life affect the individual. The so-called third wave influenced by Eastern philosophy emphasizes people's struggle to avoid unpleasant feelings and experiences as a cause of mental illness. In CBT all these perspectives are woven into the mapping of the patient's problem, and the patient with the goal of treatment to fit the patient's problems and objectives (Barlow, 2004). CBT is an active treatment for both therapist and patient. Collaboration is emphasized and the patient receives information about both their problems and the purpose of the various interventions (psychoeducation). Homework assignments are an important part of CBT, and the patient makes a large part of the work on their own between sessions.

Cognitive behavioural therapy for adolescents

Cognitive behavioural therapy has been found to be effective in treating a wide range of mental health problems in adolescents, including anxiety disorders and depression (James et al., 2015). Nonetheless, the majority of individuals with anxiety disorders do not receive evidence-based treatments (Serlachius et al., 2012; Vigerland et al., 2013). Cognitive behavioural therapy for anxiety disorders in children and adolescents often include psychoeducation, affect recognition, cognitive restructuring, relaxation, and graded exposure (Rapee et al., 2009). The treatments used may vary in the extent to which they focus on these components. The general aim, however,

is to help the patient identify physical, cognitive and behavioural components of the anxiety and then to gradually apply skills to face fearful situations outside the therapy session. Homework assignments are usually included in this process (Rapee et al., 2009) and session numbers may vary from nine to twenty sessions (James et al., 2015). Examples of evidence-based CBT treatments that are manualised for children and adolescents with anxiety disorders include Coping Cat (Kendall, 1994), Friends (Shortt et al., 2001) and Cool Kids (Hudson et al., 2009). These treatments target multiple anxiety disorders and can be administered in groups or individually.

The programs vary in the extent to which they focus on the different skills but the main goal is to help the child identify their symptoms of anxiety and use skills to gradually approach, instead of avoiding the anxiety-provoking situation. Training in other skills such as self-assertion, social skills and problem solving are also frequently included in treatment programs for children and adolescents with anxiety (Rapee, Schniering & Hudson 2009). Most CBT programs for children include parental involvement to some degree. The established evidence for treatment of anxiety disorders in children and adolescents mainly concern efficacy studies; to date, few effectiveness studies have been conducted in clinical settings (Southam-Gerow et al., 2010). In light of the developmental challenges and increased vulnerability to anxiety during adolescence, some have speculated that adolescents may be less responsive to CBT than adults (Kendall & Peterman, 2015). Adolescents' increasingly busy schedules, marked by extracurricular activities, large academic workloads, engagements, may further limit adolescents' willingness to participate in therapy and complete the necessary homework tasks.

Cognitive behavioural therapy for young adults

The first-line psychological treatment for youth anxiety disorders is cognitive behavioural therapy (Kendall & Peterman, 2015). CBT for youths has been adapted from adult protocols and addresses symptomatology that cuts across anxiety diagnoses. Specifically, CBT provides psychoeducation about anxiety, teaches youths skills for managing fears (i.e. relaxation, coping thoughts, problem solving, externalization), and provides a context for youths to gradually encounter their fears and minimize avoidance (i.e.

exposure). The core procedures in cognitive behavioural treatment of anxiety disorders in youth, defined by Woody and Ollendick (2006) and Ollendick and Hovey (2009) are cognitive restructuring, behavioural experiments and repeated exposure, and reduction of avoidance. CBT is present-focused, short-term, and active, requiring youths to participate during in-session and at-home exercises. Although CBT principles are evidence based and standardized, clinicians are encouraged to use "flexibility within fidelity," tailoring treatment to the youth's individual presentation (Kendall et al., 2008). Numerous studies have been conducted to examine CBT for anxiety disorders and anxiety symptoms in youth, and taken together, these studies provide the empirical support necessary to make CBT the only psychological treatment identified to date as an evidence-based treatment (Seligman & Ollendick, 2011). CBT for anxiety disorders in youth appears efficacious even in the presence of comorbid conditions.

Cognitive behavioural therapy for older adults

Cognitive behaviour therapy (CBT) is the most widely tested and evidencebased psychological treatment for older adults with anxiety and depression (Shrestha et al., 2011). CBT for older adults is very similar to CBT for adults at other ages, except that it is often modified to include gerontological knowledge and take account of cohort beliefs, physical health status and stage-of-life transitions (Laidlaw et al., 2008) and pleasant activities scheduling (Landreville, Gosselin, Grenier, Hudon & Lorrain, 2016). Coping strategies are included as well as techniques such as applied relaxation and mindfulness (Bohlmeijer et al., 2003). Several different forms of CBT exist but most involve consideration of maladaptive cognitions, behaviours and physiological response systems. There are some versions of CBT for older adults that have been investigated, for example problem solving therapy (Cuipers et al., 2007) and reminiscence therapy for depression (Bohlmeijer et al., 2003). More recently, an interest in acceptance-oriented CBT has emerged, but data is still sparse when it comes to older adults. Moreover, it has been argued that older adults tend to benefit somewhat less from CBT than younger adults and that "previously tested intervention protocols may not be adequately tailored to

accommodate the unique needs of older adults, and greater innovation in psychosocial interventions may be needed" (page 812, Shrestha et al., 2011).

Internet-based cognitive behavioural therapy

Therapist guided ICBT mirrors CBT but what is unique is the way the treatment is delivered to the patient and how communication is established (Andersson & Titov, 2014). ICBT require that the patient regularly logon to a secure platform where the patient over a specific time period, access, read and download online materials (i.e. modules). As in CBT the patients receive homework assignments, which they complete before the next module in their treatment. Therapist contact can take the form of real time interventions or as studied in this thesis, delayed interaction that consists of secure and encrypted e-mail communications.

CBT treatment protocols are typically designed to target one specific disorder and falls under the definition of disorder-specific CBT (Dear et al., 2015). It is, however, unclear if this is the most optimal approach in regards to the high comorbidity between anxiety disorders and depressive disorders. Internet-based CBT has in the past generally been disorder-specific and from above mentioned predicament, two alternative treatment approaches emerged, the tailored and the transdiagnostic approach that aims to simultaneously treat both principal and comorbid disorders. Previous trials on ICBT have targeted adults in general and relatively few have targeted adolescents, young adults or older adults.

ICBT for children and adolescents is a small but growing field (Vigerland et al., 2016). These groups are more heterogeneous than adults why there is a need to develop different kinds of treatment and ways to convey them becomes larger. For adolescents and young adults, ICBT has been proven effective for students with social phobia (Tillfors et al., 2011), for students with anxiety, depression and stress (Day, Wojtowicz & McGrath, 2013), for young adults with anxiety and depression (Sethi, Campbell & Ellis, 2010), for adolescents with anxiety (Spence et al., 2011) and for adolescents with obsessive-compulsive disorder (Lenhard et al., 2014).

Few trials have examined the efficacy and effectiveness of ICBT programmes for older adults with anxiety and depression. Programs for individuals aged 60 years and above have been developed in Australia, Sweden (Paper IV) and Canada. The empirical evidence for the efficacy and effectiveness of ICBT for older adults comes from a small number of trials (Dear et al., 2013; Dear et al., 2015 a, b; Titov et al., 2015; Zou et al., 2012; Staples et al., 2016; Titov et al., 2016; Jones, Hadjistavropoulos & Soucy, 2016). The results of these initial trials are, however encouraging, with moderate to large effects, and highlight the potential of ICBT as an approach of increasing access to evidence-based psychological treatment among older adults.

Tailored and transdiagnostic internet-based cognitive behavioural therapy

The transdiagnostic approach to treatment is designed to target common underlying symptoms and predisposing psychological factors for anxiety and depression (Pearl & Norton, 2017). Titov et al. (2012) has transferred this approach to internet-based CBT due to the fact that anxiety and depressive disorders share several characteristics that involve common symptoms, overall course and response to treatment. Transdiagnostic treatment offers potential advantages to disorder-specific approaches with a simplified treatment plan. Several trials have been conducted with transdiagnostic internet-based treatments with promising results (Johnston et al., 2011; Titov et al., 2013; Dear et al., 2015; Păsărelu, Andersson, Bergman Nordgren & Dobrean, 2016). Internet-based transdiagnostic CBT does not target any specific psychological disorder and is aimed to present a broad range of the apeutic information and skills to the patient (Dear et al., 2015; Fogliati et al., 2016). The information and the skills thought in this treatment format target, cognitive, physical and behavioural symptoms of psychological distress in general.

Tailored internet-based cognitive behavioural therapy is designed to identify a participants unique symptom profile and to provide information and skills that are likely to be helpful based on said profile. This identification is normally based on a diagnostic interview (i.e. SCID, ADIS) and measurements (i.e. BAI, MADRS-S) that provide the therapist with information for a behavioural analysis, that in turn guide the therapist in choosing the modules for each participant. There are exceptions to this process; in some cases it has been the participants whom have chosen the modules (Andersson, Estling, Jakobsson, Cuijpers & Carlbring, 2011). In recent years, this treatment format has been extended to also tailor on the basis of age, specifically for adolescents (Paper I), young adults (Paper II, III) and older adults (Paper IV). The modules in these trials were derived from previous studies on tailored treatment for anxiety and depression (Carlbring et al., 2011; Johansson et al., 2012; Bergman Nordgren et al., 2014). They were adapted for the different age groups both in regards to language and with clinical examples, with different fictitious case examples but sharing the same interventions. The first module (Introduction) and the last module (Relapse prevention) are fixed for all patients and then there are diagnosis and symptom specific modules that the therapist (or the patients in some trials) can choose from. The following are examples of modules that have been prescribed in the trials in this thesis: Cognitive restructuring (2) modules), Generalized anxiety (3 modules), Worrying (1 module), Social anxiety (2 modules), Panic disorder (2 modules), Panic symptoms (1 module), Agoraphobia (1 module), Applied relaxation (1 module), Behavioural activation (2 modules), Procrastination (1 module), Trauma (2 modules), Setting boundaries (1 module), Establish a better sleep pattern (1 module), Mindfulness (1 module), Problem solving (1 module) and Stress (1 module). The modules are all based on established and evidence-based CBT interventions and modules contain relevant components like psychoeducation, exposure exercises and behavioural experiments. All modules contain homework assignments for the participants, which consist of questions on the psychoeducational sections and tasks for the participant to complete. For a detailed description of the modules, see Table 1.

Table 1

Tailored internet-based treatment, a description of available modules.

Module(s)	Description		
Introduction	A brief presentation on how ICBT is		
	structured with text modules and		
	homework. The concept of anxiety is		
	introduced and different symptoms		
	and possible causes are described.		
	The basics of CBT are described and		
	information on how to set goals is		
Cognitive restructuring 1 and 2	given. These modules explain the		
cognitive restriction thig I take 2	relationship between thoughts and		
	feelings. Furthermore, the concept of		
	automatic thoughts and a model to		
	acknowledge them are presented. The		
	modules then explain how anxiety is		
	maintained via a cognitive model and		
	include strategies to challenge		
	negative automatic thoughts.		
Generalized anxiety 1, 2 and 3	The first module introduces the		
	concept of anxiety and generalized		
	anxiety, and information is given about the potential causes for concern		
	and how generalized anxiety arises.		
	In one exercise, the participants make		
	their own behavioural analysis where		
	they start from a situation with		
	concern and describe what happens		
	to the thoughts, feelings and physical		
	reactions. Thereafter, the participants		
	explain what they do when they are		
	worrying and what effect that has on		
	them.		
	In the second module on generalized		
	anxiety, strategies for reducing worrying using planned worrying and		
	worry free zones are presented. Participants may schedule their		

worrying, keep a diary of their worrying and postpone their worrying to a given time.

In the third module, participants learn to expose themselves to their worrying in order to reduce it. Participants are trained how to handle alarming situations without mentally escaping them. Participants are also encouraged to continue with scheduled worrying.

This module is a generic short version of the modules on generalized anxiety for participants who worry but do not meet the criteria for generalized anxiety disorder.

The most common symptoms, problem areas and safety behaviours in social anxiety are presented in this module. The principles of exposure are taught and information is given on how to create an anxiety hierarchy.

The second module describes a cognitive model of social anxiety with the aim of providing a thorough knowledge of what maintains maladaptive behaviours. In various exercises, participants practice shifting focus from themselves to the surroundings in order to become less anxious and self-conscious.

These modules provide psychoeducation on panic disorder. Participants are then guided in interoceptive exposure.

This module provides participants with the knowledge of what panic attacks are and how panic attacks are linked to breathing. In this module, the participant implements various

Worrying

Social anxiety 1 and 2

Panic 1 and 2

Panic symptoms

Agoraphobia

Applied relaxation

Behavioural activation 1 and 2

Procrastination

breathing tests to investigate if this provokes panic symptoms and the participants get knowledge of hyperventilation during panic attacks and how to stop this by regulated breathing.

In this module the participant is presented with information about agoraphobia and the consequences of avoiding different situations. Types of safety behaviours are described and their implications. Exposure guidelines are presented as well as how to design goals for exposure.

This module introduces how to use applied relaxation to reduce anxiety/worrying. The module contains an audio file. Participants are encouraged to practice relaxation daily and continuously evaluate their effects.

These modules begin by describing depression, its causes and consequences from a CBT perspective. The aim of behavioural activation is explained. Participants list preferred behaviours and register them in their activity plan.

In the second module of behavioural activation, participants continue the development of their activity plan. They receive information on how they can reward themselves for the implementation of activities and they are encouraged to evaluate and develop their activity plan.

The module provides psychoeducation about behavioural patterns of postponing activities or tasks. Strategies for handling procrastination are then introduced.

Trauma 1 and 2

given psychoeducation about posttraumatic stress symptoms and common reactions after a trauma. The participants are then guided in exposure exercises and the module provides information about common problems that may arise during exposure.

In the first module, participants are

In the second module, participants use cognitive restructuring as a way to relate different to their thoughts about their trauma.

This module is aimed towards participants who have difficulty setting boundaries. Advice on setting boundaries and concrete techniques for saying 'no' are presented in the module. Furthermore, the module focuses on relationships and how to create and maintain good relationships while maintaining self-respect.

In this module, the participant is given psychoeducation about sleep and sleep hygiene. The participant must then identify their sleep pattern and examine the factors that influence their sleep patterns and what needs to change. Different sleep strategies are presented, and participants are encouraged to test the strategies that seem appropriate based on their own sleep patterns.

In this module, participants are introduced to mindfulness and the module consists of several mindfulness exercises. The exercises focus on different areas: the breathing, the body and emotions. For each exercise, there are audio files that can be used as guidance.

Setting boundaries

Establish a better sleep pattern

Mindfulness

Problem solving

Stress

Relapse prevention

In this module, the participants practice problem-solving skills as a means of coping with stressful problems.

In this module, the participants receive psychoeducation about stress. The participant then identifies situations that cause them stress. The module highlights the importance of a balance between rest and activity and sets an action plan to accomplish this. The module also has a section with work related stress and concrete advice is given on how stress can be reduced.

The final module describes the difference between setbacks and relapse. Common risk situations for setbacks are described as well as general strategies for dealing with these. Participants are encouraged to summarize their treatment where they are asked to describe what was most important to them based on modules they have gone through. Participants are also invited to go through what they have learned and make a written plan of action for the future.

Setting

Guided ICBT has been investigated in more than 100 controlled efficacy studies (Hedman et al., 2012). In these efficacy studies internal validity is prioritized, which means that experimental control is important. In practice, this means that research participants tend to be highly selected, homogenous in clinical characteristics, self-referred, and therapists are well trained and monitored for adherence to treatment manuals (Andersson & Hedman, 2013). Moreover, efficacy studies tend to be conducted at research university clinics where therapists usually have smaller caseloads sometimes supported by students in training. In effect, it is often unclear if

the results of such studies can be generalized to routine practice. An effectiveness study examines whether a treatment works in clinical settings and in situations that clinicians encounter in their daily routine practice (Lutz, 2003). It can be argued however that clinical intervention research and trials on psychotherapy, always have elements of effectiveness as they include real patients with disorders that are indeed both comorbid and severe (Stirman et al., 2003). The support for the efficacy of guided ICBT for mood and anxiety disorders has been covered in several systematic reviews and meta-analyses with effect sizes similar to those of regular CBT (Spek et al., 2007; Andrews et al., 2010; Hedman et al., 2012). Andersson and Hedman (2013) defines effectiveness studies as a study conducted in a setting equivalent or similar to routine clinical care where patients are not solely recruited through self-referral and treatments are delivered by staff with permanent employment. The efficacy of ICBT has been proven in a vast number of trials and the number of trials investigating the effectiveness of ICBT is increasing but is however, few in number (Andersson & Hedman, 2013; Andersson & Titov, 2014). The increasing number of effectiveness studies on ICBT for anxiety and mood disorders includes studies on panic disorder, social phobia, generalized anxiety disorder, posttraumatic stress disorder and depression (Hedman et al., 2013). These trials suggest that the effects found in controlled efficacy trials on guided ICBT tend to be replicated in clinical practice.

While ICBT is a promising treatment option that could be used to increase accessibility to evidence-based psychological treatment, it is of importance to investigate whether it is also effective when delivered in a routine care setting. Without effectiveness studies there is a risk of implementing new treatments that might not work in contexts outside the highly controlled randomised trials that have been conducted. As effectiveness studies on ICBT for different age groups are lacking it is important to increase the body of knowledge in this area by investigating how the treatment works in a routine care setting. From the existing effectiveness trials it is indicated that it is possible to transfer ICBT to clinical practice with sustained effects and moderate to large effect sizes (Andersson & Hedman, 2013). In this thesis both efficacy studies (Paper I and IV) and effectiveness studies (Paper II and III) are included.

Overall aim

The overall aim of this thesis is to further develop and test the effects of tailored internet-based cognitive behavioural therapy on the basis of age, for adolescents, young adults and older adults. Specifically by:

Developing and testing the effects of individually tailored internet-based cognitive behavioural therapy for adolescents with anxiety and comorbid depressive symptoms within a child and adolescent psychiatric clinic.

Adapting and testing the effects of individually tailored internet-based cognitive behavioural therapy for young adults and adults with panic attacks with comorbid anxiety and depressive symptoms in a randomized controlled trial.

Adapting and testing the effects of individually tailored internet-based cognitive behavioural therapy for young adults with anxiety and comorbid depressive symptoms within a Youth Health Care Centre.

Adapting and testing the effects of individually tailored internet-based cognitive behavioural therapy for older adults with anxiety and comorbid depressive symptoms in a randomized controlled trial.

Empirical studies

Paper I. Silfvernagel, K., Gren-Landell, M., Emanuelsson, M., Carlbring, P., & Andersson, G. (2015). Individually tailored internet-based cognitive behavior therapy for adolescents with anxiety disorders: A pilot effectiveness study. *Internet Interventions*, *2*, 297-302.

Aim

The aim of Paper I was to develop and test the effects of individually tailored ICBT for adolescents with anxiety and comorbid depressive symptoms. The intervention was developed and tested within a child and adolescent psychiatric clinic.

Methods

In this single-group open trial, participants were recruited through ordinary referral routes (i.e. self-referral, referral through a parent/guardian/doctor) within a child and adolescent psychiatric clinic. Information about the study was given on a website hosted by the clinic, and brochures and information sessions were offered to all student counseling facilities in the area. After being referred to the trial, participants met with the project leader for an initial assessment to see if they met the inclusion criteria and to receive further information about the project. During this initial meeting written consent was obtained. Inclusion criteria for the trial required that the participant be 15-19 years old, or turning 15 within the year of entering the trial, and seeking treatment at the clinic for mild to moderate anxiety, including: social anxiety, excessive worrying or panic attacks. Exclusion criteria were that the participant met diagnostic criteria for obsessivecompulsive disorder, post-traumatic stress disorder, severe depression, risk of suicide, suffering from a severe psychiatric disorder (i.e. bipolar disorder and psychosis), had ongoing alcohol abuse/dependence, or other social or psychological problems deemed to overshadow the anxiety problems. Participants could not be in psychological treatment and if on medication had to be on a stable dose for at least three months. Participants who met the inclusion criteria were asked to complete an online screening via the internet treatment platform. In addition to background questions (i.e. questions regarding the adolescents living situation, number of siblings, occupation of their parents and the family's economical situation, see Table 2), this included the Beck Anxiety Inventory (BAI; Beck et al., 1988), Clinical Outcomes in Routine Evaluation-Outcome Measure (CORE-OM; Barkham et al., 2001), Montgomery-Åsberg Depression Rating Scale-Self-Rated (MADRS-S; Svanborg & Åsberg, 1994) and the Alcohol Use Disorders Identification Test (AUDIT; Saunders et al., 1993).

The semi-structured interview, Anxiety Disorders Interview Schedule for DSM-IV: Child and Parent Versions (ADIS-C/P; Silverman & Albano, 1996) was administrated face-to-face to determine present diagnosis for the adolescent along with providing information for a behavior-analysis to determine treatment prescription. The adolescent was not required to meet the diagnostic threshold for an anxiety disorder on the ADIS-C/P to participate in the study. At post-treatment the same procedure was repeated with the exception of AUDIT, which was only used at screening.

Treatment consisted of tailored ICBT adapted for adolescents. The duration of treatment was 6-18 weeks with a prescription of 6-9 treatment modules for each participant. The participants were advised to spend 1–2 weeks on each prescribed module and the treatment was combined with additional telephone calls or face-to-face sessions if required. Treatment consisted of 17 potential modules, with different fictitious case examples. The language was adapted to suit the age group. The modules were derived from previous studies on tailored ICBT for anxiety and depression (Andersson et al., 2011; Carlbring et al., 2011; Johansson et al., 2012; Bergman Nordgren et al., 2014; Silfvernagel et al., 2012). All modules were based on established and evidence-based CBT principles (i.e. exposure). The modules included psychoeducation, exposure exercises, behavioral experiments homework assignments. Two psychologists provided therapist guidance. The treatment was individually tailored for each participant based on the results of the ADIS-C/P interview. The ADIS-C/P interview provided the therapist with information for a behavioral analysis that in turn guided the therapist in choosing the modules for each adolescent. For example if the adolescent presented symptoms of social anxiety along with excessive

worrying they could receive the modules Introduction, Cognitive restructuring 1 and 2, Social anxiety 1 and 2, Worrying and Relapse prevention.

Changes in questionnaire scores between pre- and post-treatment were evaluated using paired sample *t*-tests. The within-group effect sizes, Cohen's *d*, was calculated from the observed means and observed pooled standard deviations from pre- to post-measurement. Non-response was handled based on the principle Complete Cases (Salim, MacKinnon, Christensen & Griffiths, 2008), which means that only data from the participants who completed the survey were included in the analysis.

Table 2. Demographic description of the participants at pre-treatment.

	Treatment group $(n = 11)$
Gender, n (%)	
Male	5 (45.5%)
Female	6 (54.5%)
Age (years)	
Mean (SD)	16.8 (1.66)
Minimum-maximum	14-19
Living situation, n (%)	
With mom and dad	8 (72.7%)
With mom	2 (18.2%)
Other	1 (9.1%)
Siblings, n (%)	
Siblings	11 (100%)
Younger siblings	5 (45.5%)
Older siblings	3 (27.3%)
Younger and older siblings	3 (27.3%)
Financial family situation, n (%)	
Very good	5 (45.5%)
Pretty good	3 (27.3%)
Some financial difficulties	3 (27.3%)
Employment status of parents, n (%)	
Both employed	9 (81.8%)
One on sick leave/ unemployed and one	2 (18.2%)
employed	
Diagnosis based on ADIS, n (%)	_
Generalized anxiety disorder	8 (72.7%)
Panic disorder	4 (36.4%)
Panic disorder + agoraphobia	1 (9.1%)
Social phobia	7 (63.6%)
Specific phobia	5 (45.5%)
Dysthymia	2 (18.2%)
Comorbid disorders	7 (63.6%)

ADIS = Anxiety Disorders Interview Schedule for DSM-IV

Results and discussion

Paired sample *t*-tests showed significant results across all outcome measures, primary and secondary, with large within-group effect sizes. At post-treatment, 5 of 8 participated in the ADIS interview. Out of these adolescents 80 percent (4/5) no longer met DSM-IV criteria for their primary anxiety disorder. Of the 11 adolescents allocated to treatment, 8 completed a mean of 6.5 modules out of 6-9 modules prescribed. Six adolescents chose to use the option with additional sessions face-to-face at the clinic and 6 adolescents asked for additional telephone-based consultations during treatment. Three of the parents asked for parental support during the treatment.

The significant effects that were found on all dependent measures immediately following treatment and adherence rates for the 8 adolescents are consistent with previous trials on CBT and ICBT for adolescents (Spence et al., 2011; Tillfors et al., 2011; Lenhard et al., 2014), and with previous trials on tailored ICBT for young adults and adults (Carlbring et al., 2011; Paper II). Overall, the results found in this study suggest that tailored ICBT may be a suitable treatment for adolescents with anxiety. Existing studies, including this study, indicate promising results but larger studies are needed. There is also limited empirical data to date that clearly indicates whether and when adolescents might require or benefit from ICBT programs designed specifically for their age group. The significant results and large within-group effect sizes reported in this study should be viewed with caution given the most important limitation of this study: the small sample size and the lack of randomization to a control group, limiting internal validity of the study. In addition, there was no follow-up of treatment results after the post-assessment. However, the study was conducted in a clinical setting with a population that is common in child and adolescent psychiatry.

Paper II. Silfvernagel, K., Carlbring, P., Kabo, J., Edström, S., Eriksson, J., Månson, L., & Andersson, G. (2012). Individually tailored internet-based treatment for young adults and adults with panic attacks: Randomized controlled trial. *Journal of Medical Internet Research*, 14 (3), e65.

Aim

The aim of Paper II was to test the effects of individually tailored ICBT for young adults and adults with panic attacks with comorbid anxiety and depressive symptoms in a randomized controlled trial.

Methods

The participants were recruited via an online list among individuals who had expressed an interest in participating in research on internet-based cognitive behavior therapy for panic disorder and generalized anxiety disorder. They were presented with the project platform, which contained information about the trial, how to register, and how to submit written informed consent. Screening consisted of the following questionnaires via the internet: Montgomery-Åsberg Depression Scale-Self-rated (MADRS-S; Svanborg & Åsberg, 1994); Clinical Outcomes in Routine Evaluation-Outcome Measure (CORE-OM; Barkham et al., 2001); Beck Anxiety Inventory (BAI; Beck, Epstein, Brown & Steer, 1988); Quality of Life Inventory (QOLI; Frisch, Cornell, Villanueva & Retzlaff, 1992); Alcohol Use Disorders Identification Test (AUDIT; Saunders, Aasland, Babor, de la Fuente & Grant, 1993); and 13 additional questions with reference to demographic variables. If the participants met the initial inclusion criteria they underwent further screening consisting of the Structured Clinical Interview for DSM-IV Axis I Disorders (SCID-I; First, Spitzer, Gibbon & Williams, 2002) and the primary outcome measure, the Panic Disorder Severity Scale (PDSS; Shear et al., 1997), conducted over the telephone by three clinical psychology MSc students who had completed their clinical training. The participants had to have reoccurring panic attacks to be included. They could also fulfill the Diagnostic and Statistical Manual of Mental Disorders, 4th edition, text revision (APA, 2000) criteria for any specific anxiety disorder, or anxiety disorder not otherwise specified except for obsessive-compulsive disorder and posttraumatic stress disorder, which would lead to exclusion. Participants could also meet the criteria for comorbid major depression, but not as the primary disorder. The participants had to be between the ages of 18 and 30 years or between 31 and 45 years; have a total score of <31 on the MADRS-S and a score <4 points on item 9 (suicidal thoughts) on MADRS-S; not currently be in psychotherapy; if on medication, be on stable dosage for the last 3 months; and not be at risk of alcohol abuse or fulfilling the criteria for current alcohol addiction. 149 individuals expressed an interest in the trial, which commenced in February 2010. After screening and diagnostic interview, 57 participants were included. For a demographic description of the participants, see Table 3. The participants were divided into two groups so that the two predetermined age groups 18-30 years (young adults) and 31–45 years (adults) were equally represented in each condition. The blocked randomization process was conducted through an online true random number-generation service (random.org) independent of the investigators and therapists. At post treatment participants were instructed via email to complete the follow-up questionnaires and to participate in a semi-structured telephone interview carried out by a blinded assessor who had no earlier contact with the participants. The same procedure was repeated at 12 months after treatment completion.

The treatment consisted of 19 CBT modules derived from previous ICBT trials on panic disorder (Carlbring et al., 2006), generalized anxiety disorder, social phobia (Andersson et al., 2006), depression (Vernmark et al., 2010), and tailored ICBT for anxiety and depression (Carlbring et al., 2011). In this trial the first module (introduction) and the last module (relapse prevention) were fixed, and the following 17 were optional for the therapists to prescribe: cognitive restructuring (2 modules); panic disorder (2 modules); agoraphobia (1 module); generalized anxiety (3 modules); social anxiety (2 modules); behavioral activation (2 modules); applied relaxation (1 module); stress (1 module); mindfulness (1 module); problem solving (1 module); and establish a better sleep pattern (1 module). The modules are all based on established and evidence-based CBT components. The panic modules, for example, consisted of psychoeducation and interoceptive exposure. All modules included psychoeducation, nearly all contained exposure exercises,

and some contained behavioral experiments depending on the content. All modules contained homework assignments for the participants, which consisted of questions on the psychoeducational sections and tasks for the participant to complete, such as exposure exercises. The aim was to prescribe 6–8 modules within an 8-week time frame for each participant. A typical prescription for the participants could be an introduction, cognitive restructuring 1 and 2, panic disorder 1 and 2, agoraphobia, applied relaxation, and relapse prevention. Therapist guidance was included in the trial. The therapists were three clinical psychology MSc students who had completed their clinical training and who were supervised by experienced clinical psychologists (senior authors). The therapists were responsible for 9–10 participants each during the 8 weeks of treatment. The treatment was individually tailored for each participant based on the results of the SCID-I interview and the clinical impression from the telephone interview. The participants were required to have access to a computer with an Internet connection and be able to download the prescribed modules in PDF format through an encrypted contact system, which they also used when communicating with their therapist. The participants were advised to spend 1 week on each prescribed module. The therapists spent approximately 15 minutes per week with each participant (estimated). The control condition consisted of a waitlist group. Participants were informed that they would receive the treatment after 10 weeks, when the treatment group had completed their treatment.

A mixed-models approach with an unstructured covariance structure was endorsed as a way to handle missing data at post treatment and at the 1-year follow-up. As suggested by Gueorguieva and Krystal (2004), the mixed-effect models approach was used due to their advantages over traditional methods of repeated-measures analysis. The between-group and within-group effect sizes (Cohen's d) was calculated from estimated means and observed pooled standard deviations. To examine whether the randomization process had succeeded in generating a balanced distribution across the two conditions, independent t-tests and a chi-square test were used for the demographic data and pre-treatment measures.

 Table 3. Demographic description of the participants at pre-treatment.

		Treatment $(n = 29)$	Control $(n = 28)$	Total $(n = 57)$
Gender	Male	8 (27.6%)	12 (42.9%)	20 (35.1%)
	Female	21 (72.4%)	16 (57.1%)	37 (64.9%)
Age	Mean (SD)	32.3 (7.4)	32.5 (6.5)	32.4 (6.9)
	Min-Max	20-45	21-44	20-45
	18-30	13 (44.8%)	12 (42.9%)	25 (43.9%)
	31-45	16 (55.2%)	16 (57.1%)	32 (56.1%)
Highest	Nine year	2 (6.9%)	2 (7.1%)	4 (7.0%)
educational	compulsory			
level	school	11 (27 00()	1 (1 1 20 ()	15 (26 22 ()
	Secondary school	11 (37.9%)	4 (14.3%)	15 (26.3%)
	Vocational	1 (3.4%)	3 (10.7%)	4 (7.0%)
	College/univ	5 (17.2%)	7 (25.0%)	12 (21.1%)
	ersity (not			
	compl.)	10 (24 50()	10 (10 00()	22 (22 (24)
	College/univ ersity	10 (34.5%)	12 (42.9%)	22 (38.6%)
	(compl.)			
Psychotherapy	No	10 (34.5%)	6 (21.4%)	16 (28.1%)
1 Sychotherapy	experience	10 (34.370)	0 (21.470)	10 (28.170)
	Previous	19 (65.5%)	22 (78.6%)	41 (71.9%)
	experience			
Anxiolytic	Ongoing	15 (51.7%)	12 (42.9%)	27 (47.4%)
and/or	Completed	4 (13.8%)	5 (17.9%)	9 (15.8%)
antidepressant	No	10 (34.5%)	11 (39.3%)	21 (36.8%)
medication	experience			
Employment	Self-	2 (6.9%)	0 (0%)	2 (3.5%)
status	employed			
	Employed	14 (48.3%)	14 (50.0%)	28 (49.1%)
	Unemployed	0 (0%)	4 (14.3%)	4 (7.0%)
	Student	5 (17.2%)	6 (21.4%)	11 (19.3%)
	On parental	4 (13.8%)	2 (7.1%)	6 (10.5%)
	leave	4 (12 00/)	0 (7 10/)	((10 50/)
	Sick leave	4 (13.8%)	2 (7.1%)	6 (10.5%)

SCID-diagnosis	Panic disorder	2 (6.9%)	2 (7.1%)	4 (7.0%)
	Panic disorder + agoraphobia	24 (82.8%)	23 (82.1%)	47 (82.5%)
	Generalized anxiety disorder	2 (6.9%)	9 (32.1%)	11 (19.3%)
	Social phobia	1 (3.4%)	8 (28.6%)	9 (15.8%)
	Anxiety disorder not otherwise specified	1 (3.4%)	0 (0%)	1 (1.8%)
	Major depression	2 (6.9%)	3 (10.7%)	5 (8.8%)
	Suffering from comorbid	5 (17.2%)	13 (46.4%)	18 (31.6%)
GCID G:	disorders	· C DGI	(III.4 · I.D.	7

SCID = Structured Clinical Interview for DSM-IV Axis I Disorders

Results and discussion

The treatment was superior to the control condition with significant interactions on all measures. A mixed-models analysis of the immediate results of the primary outcome measure and the secondary outcome measures, showed a significant interaction and moderate to large between-group effect sizes were observed. Age group did not interact with treatment condition across any measure. There was, however, an interaction of time and age group for BAI. Of the 29 participants in the treatment group, 7 (24%) completed all prescribed modules (6–8) within the 8-week treatment period. A total of 17 participants (59%) completed 50 percent of the prescribed modules and 14 (48%) completed 75 percent of the prescribed modules. The mean number of completed modules for the whole group was 5.0 (SD 2.6). The mean number of completed modules in the young adult group was 5.15 (SD 2.34) and the corresponding number in the adult group was 4.19 (SD 3.16). This difference was not statistically significant. At the

1-year follow-up a mixed-models analysis showed significant time effects for the primary outcome measure and for the secondary outcome measures.

The results of this trial, with moderate to large effect sizes, are consistent with previous trials of ICBT for panic disorder (Carlbring et al., 2006) and transdiagnostic ICBT treatment for anxiety (Titov, Andrews, Johnston, Robinson & Spence, 2010). The results of his trial, although focused on individuals with reoccurring panic attacks, are hence in line with previous trials of tailored ICBT for anxiety (Carlbring et al., 2011; Păsărelu et al., 2016). No interaction effect between age group was found; there was, however, an interaction between time and age group for the BAI. This could indicate that the young adults as a group are more likely to improve in the short time frame. The effect sizes across all measures showed a tendency for larger effects among the young adults but due to the small sample size, this requires further investigation for any conclusions to be drawn.

Paper III. Silfvernagel, K., Wassermann, C., & Andersson, G. (accepted). Individually tailored internet-based cognitive behavioural therapy for young adults with anxiety disorders: A pilot effectiveness study. *Internet Interventions*.

Aim

The aim of Paper III was to determine whether individually tailored ICBT is a feasible approach in the treatment of anxiety symptoms and comorbid anxiety and depressive symptoms for young adults in a clinical setting.

Methods

This interventional study with a single group assignment took place in a Youth Health Care Centre in Sweden. The study was approved by the regional ethics committee in Linköping and registered at ClinicalTrials.gov (NCT01402258). Participants were recruited from the Centre via selfreferral through an early version of the online platform developed by the research group (Vlaescu, Alasjö, Miloff, Carlbring & Andersson, 2016). The platform contained information about the trial, how to register, and how to submit written informed consent. Screening consisted of the following questionnaires via the Internet: Beck Anxiety Inventory (BAI; Beck et al., 1988) (primary outcome measure); Montgomery-Asberg Depression Scale-Self-rated (MADRS-S; Svanborg & Åsberg, 1994); Clinical Outcomes in Routine Evaluation-Outcome Measure (CORE-OM; Barkham et al., 2001); Quality of Life Inventory (QOLI; Frisch, Cornell, Villanueva & Retzlaff, 1992); Alcohol Use Disorders Identification Test (AUDIT; Saunders, Aasland, Babor, de la Fuente & Grant, 1993); and additional questions with reference to demographic variables. If the participants met the initial inclusion criteria they underwent further screening consisting of the Structured Clinical Interview for DSM-IV Axis I Disorders (SCID-I; First, Spitzer, Gibbon & Williams, 2002), conducted by licensed psychologists. The licensed psychologists then presented their clinical assessment to the main author and a psychiatrist before the participants were enrolled in the trial. This procedure was then repeated post treatment with the exception of AUDIT. The participants had to be between the ages of 16-25 years old, young adults, and have reoccurring anxiety symptoms to be included. They

could also fulfill the Diagnostic and Statistical Manual of Mental Disorders, 4th edition, text revision criteria for any specific anxiety disorder, or anxiety disorder not otherwise specified. Participants could also meet the criteria for comorbid major depression, but not as the primary disorder. The participants had have a score of <4 points on item 9 (suicidal thoughts) on MADRS-S; not currently be in psychotherapy; and not be at risk of alcohol abuse or fulfilling the criteria for current alcohol addiction. 55 individuals expressed an interest in the trial, of which 40 filled in all the self-assessment forms. 11 were referred to child and adolescent psychiatry after this initial screening and 9 declined to participate. 20 were assessed with SCID-I, 5 individuals were referred to child and adolescent psychiatry and psychiatry and 15 participants were included the study. For a demographic description of the participants, see Table 4.

The experimental treatment in this trial consisted of a behavioural intervention that consisted of tailored Internet-administered CBT. The treatment consisted of 22 modules with accompanying homework assignments. The modules were between 9 and 45 pages long, with an average length of 19.7 pages. The modules were derived from previous studies on tailored treatment for anxiety and depression (Carlbring et al., 2011; Silfvernagel et al., 2012). The modules were adapted for young adults both in regards to language and with clinical examples. The modules are all based on established and evidence-based CBT principles and included psychoeducation, exposure exercises, behavioural experiments and homework assignments. The treatment was individually tailored for each participant based on the results of the clinical assessment based on the questioners and the SCID-I interview. The first module (introduction) and the last module (relapse prevention) were fixed, and the following were available for the psychologist to prescribe: cognitive restructuring (2) modules); panic disorder (2 modules); panic symptoms (1 module); agoraphobia (1 module); social anxiety (2 modules); trauma (2 modules); setting boundaries (1 module); behavioural activation (2 modules); procrastination (1 module); worrying (1 module); generalized anxiety (3 modules); applied relaxation (1 module); and establish a better sleep pattern (1 module). Licenced and experienced psychologists provided therapist guidance and there were no automatic messages.

Changes in questionnaire scores between pre- and post-treatment were evaluated using paired sample t-tests. The within-group effect sizes (Cohen's d) were calculated from the observed means and observed pooled standard deviations from pre- to post-measurement. Non-response was handled based on the principle Complete Cases (Salim et al., 2008), which means that only data from the participants who completed the survey were included in the analysis. Also examined was the percentage of participants who no longer met a diagnosis after treatment.

Table 4. Demographic description of the participants at pre-treatment.

	Treatment group $(n = 15)$
Gender, n (%)	
Male	1 (6.7%)
Female	14 (93.3%)
Age (years)	
Mean (SD)	21.5 (1.8)
Minimum-maximum	18–24
Marital status, n (%)	
Single	7 (46.7%)
Living together with a partner	6 (40%)
Living with parents	2 (13.3%)
Highest educational level, n (%)	
Secondary school	10 (66.7%)
College/university (not completed)	5 (33.3%)
Psychotherapy, n (%)	
No experience	7 (46.7%)
Previous experience	8 (53.3%)
Anxiolytic and/or antidepressant, n (%)	((((((((((((((((((((
Ongoing	3 (20%)
Completed	2 (13.3%)
No experience	10 (66.7%)
Employment status, n (%)	
Employed	6 (40%)
Unemployed	1 (6.7%)
Student	7 (46.7%)
Sick leave	1 (6.7%)
SCID-I diagnosis, n (%)	
Generalized anxiety disorder	4 (26.7%)
Panic disorder	3 (20%)
Panic disorder + agoraphobia	5 (33.3%)
Social phobia	1 (6.7%)
Obsessive-compulsive disorder	1 (6.7%)
Posttraumatic stress disorder	3 (20%)
Anxiety disorder not otherwise	1 (6.7%)
specified	
Major depression	6 (40%)

SCID-I = Structured Clinical Interview for DSM-IV Axis I Disorders

Results and discussion

Paired sample *t*-tests showed significant results across all outcome measures, primary and secondary, with large within-group effect sizes. According to the assessment of the SCID interview after treatment 7 of 10 participants no longer met the criteria for a diagnosis. Of the 15 participants who were included in the study, 9 completed all prescribed modules (8-12) with a mean of 10.1 modules. One participant announced after five of the eight planned modules that they desired to end treatment because they were feeling much improved and thereafter participated in post-measurement. The 5 who discontinued treatment prematurely finished a mean of 3.4 modules and did not participate in the post-measurement assessment.

The results of this study are consistent with study one of individually tailored ICBT conducted in a clinical setting for adolescents. The results are also in line with Halje et al. (2015) who examined CBT treatment at youth health care Centers within the same county where this study took place. The results should be viewed with caution due to a small sample size and the lack of randomization, however this was a pilot effectiveness study with experienced psychologists that conducted the interviews and administrated the treatment that strengthens the validity. An essential aspect is how well the sample in the study represents the population studied. The treatment was tested in regular care with a patient population that is common and representative for young adults seeking first-line treatment. Inclusion criteria were low with few restrictions in terms of comorbidity, which mimics a clinical population, which enhances the external validity for this study.

Paper IV. **Silfvernagel, K.**, Westlinder, A., Andersson, S., Bergman, K., Diaz Hernandez, R., Fallhagen, L., Lundqvist, I., Masri, N., Viberg, L., Forsberg, M-L., Lind, M., Berger, T., Carlbring, P., & Andersson, G. (submitted). Individually tailored internet-based cognitive behaviour therapy for older adults with anxiety and depression: A randomised controlled trial.

Aim

The overall aim of Paper IV was in a randomised controlled trial to test the effects of individually tailored ICBT for anxiety and depression in order to generate an evidence-based treatment alternative that could increase access to psychological treatments for older adults. The second aim was to investigate if pre-treatment cognitive flexibility and self-perceived cognitive functioning would predict treatment outcome.

Methods

In this two-group RCT the participants were recruited via advertising in the Daily Mail. Seventy-nine individuals were assessed for eligibility and sixtysix participants were included in the trial and randomised to the two conditions individually tailored ICBT or general support from a therapist. Table 5 provides a summary of participant characteristics. The assessment consisted of the self-report questionnaires Beck Anxiety Inventory (BAI; Beck et al., 1988), Generalized Anxiety Disorder 7-item scale (GAD-7; Spitzer et al., 2006), Montgomery Åsberg Depression Rating Scale - Self Rated (MADRS-S; Svanborg & Asberg, 1994), Patient Health Questionnaire-9 (PHQ-9; Kroenke et al., 2001), Clinical Outcomes in Routine Evaluation-Outcome Measure (CORE-OM; Evans et al., 2000), Quality of Life Inventory (QOLI; Frisch et al., 1992) and the Alcohol Use Disorders Identification Test (AUDIT; Saunders, Aasland, Babor, De la Fuente & Grant, 1993). The Structured Clinical Interview for DSM-IV Axis I Disorders (SCID-I; First, Gibbon, Spitzer, & Williams, 2002) was administered to establish diagnoses for the participants and to tailor the treatment. In order to investigate if pre-treatment cognitive flexibility and self-perceived cognitive functioning would predict treatment outcome an

online in-house developed version of the standard, single-deck version of the Wisconsin Card Sorting Test-64 (WCST; Greve, 2001) featuring 64 response cards was included in the pre-treatment assessment along with the Cognitive Failure Questionnaire (CFQ; Broadbent et al., 1982) that measurers self-perceived cognitive functioning. The same procedure was then conducted at post-treatment.

The treatment group received individually tailored ICBT for symptoms of anxiety and depression while the control group received general support from a therapist. The control group was offered the same treatment after the initial trial period, that is, after the treatment group had finished the 8-week treatment. The individually tailored treatment is designed to identify participant's unique symptom profile and to provide information and skills that were likely to be helpful based on a participant's symptom profile. Individually tailored ICBT aims to tailor the treatment according to the participant's needs and symptoms with transdiagnostic components (Silfvernagel et al., 2012). The goal is to identify and target participant's specific psychosocial difficulties and psychological comorbidities with the aim of increasing the relevance, comprehensiveness and clinical outcomes of ICBT treatments. Importantly, the treatment package for older adults consists of modules derived from previous ICBT treatments (Carlbring et al., 2011; Andersson et al., 2011; Johansson et al., 2012; Silfvernagel et al., 2012; Bergman Nordgren et al., 2013), which have been adapted to make them suitable for an older population. The first module (the introduction module) and the last module (a relapse prevention module) are fixed and the following are optional for the therapists to prescribe within an 8-week timeframe: cognitive restructuring (2 modules), panic disorder (2 modules), agoraphobia (1 module), generalised anxiety (3 modules), social anxiety (2 modules), behavioural activation (2 modules), applied relaxation (1 module), stress (1 module), mindfulness (1 module), problem-solving (1 module) and sleep habits (1 module). There are also long and short versions for the diagnosis-specific modules for the therapist to choose from. The modules are all based on established and evidence-based CBT treatments and modules contain relevant components like psychoeducation, exposure exercises and behavioural experiments. All modules contain homework assignments for the participants, which consist of questions on the

psychoeducational sections and tasks for the participant to complete, such as exposure exercises. Therapist guidance is included in the treatment and there are no automatic emails; either the therapist or the participant initiates all contact.

A mixed-model approach with an unstructured covariance structure was endorsed to examine treatment effects immediately after treatment for the main and secondary outcome measures with the assumption of data missing at random (MAR) for full maximum likelihood estimation (ML). With this approach an unbiased estimate of the average casual effect is obtained including all the individuals included in the trial (Hesser, 2015). Betweengroup effect sizes (Cohen's d) were calculated from estimated means and observed pooled standard deviations. To examine whether the randomization process had succeeded in generating a balanced distribution across the two conditions, independent t-tests were used for the demographic data and pre-treatment measures. To examine whether cognitive flexibility measured by perseverative errors on the WCST and initial levels of self-perceived cognitive functioning measured by the CFQ predicted outcome we calculated change scores and used pre-treatment scores as covariates

Table 5. Demographic description of the participants at pre-treatment.

		Treatment group $(n = 33)$	Control group $(n = 33)$	Total $(n = 66)$
Gender	Female	22 (66.7%)	28 (84.8%)	50 (75.8%)
Age (years)	Male Mean Minimum– maximum	11 (33.3%) 66.7 60 – 77	5 (15.2%) 65.5 60 – 73	16 (24.2%) 66.1 60-77
Marital status	Single Married/ living together	11 (33.3%) 17 (51.5%)	10 (30.3%) 18 (54.5%)	21 (31.8%) 35 (53%)
	Living apart	3 (9.1%)	3 (9.1%)	6 (9.1%)
	Widow/ widower	2 (6.1%)	2 (6.1%)	4 (6.1%)
Highest educational level	Compulsory school	4 (12.1%)	3 (9.1%)	7 (10.6%)
	Secondary school	6 (18.2%)	8 (24.2%)	14 (21.2%)
	Vocational College/ university	7 (21.2%) 16 (48.5%)	3 (9.1%) 19 (57.6%)	10 (15.2%) 35 (53%)
Employment status	Retired	26 (78.8%)	22 (66.7%)	48 (72.7%)
	Employed Sick leave	3 (9.1%) 1 (3.0%)	10 (30.3%) 0 (0%)	13 (19.7%) 1 (1.5%)
Psycho- therapy	Unemployed No experience	3 (9.1%) 8 (24.2%)	1 (3.0%) 13 (46%)	4 (6.1%) 21 (34.6%)
Anxiolytic	On-going	15 (45.5%)	14 (42.4%)	29 (43.9%)
and/or antidepressant medication	Completed No experience	12 (36.4%) 6 (18.2%)	10 (30.3%) 9 (27.3%)	22 (33.3%) 15 (22.7%)
Somatic illness	Self-reported	19 (57.6%)	18 (54.5%)	37 (56.1%)
Medication for somatic illness	On-going	17 (51.5%)	18 (54.5%)	35 (53%)

Psychiatric diagnosis	Generalized anxiety disorder	19 (57.6%)	16 (48.5%)	35 (53%)
	Panic disorder	5 (15.2%)	6 (18.2%)	11 (16.7%)
	Panic disorder with agoraphobia	4 (12.1%)	0 (0%)	4 (6.1%)
	Social phobia	4 (12.1%)	2 (6.1%)	6 (9.1%)
	Posttraumatic stress disorder	2 (6.1%)	2 (6.1%)	4 (6.1%)
	Major depression	13 (39.4%)	8 (24.2%)	21 (31.8%)

Results and discussion

The mixed-model analysis of the immediate results of the primary outcome measure of anxiety showed a significant interaction effect. For the secondary outcome measures of anxiety, depression and quality of life, significant interaction effects were observed with moderate to large between-group effect sizes with the exception of quality of life wehre a small between-group effect size was observed. Of the 33 participants randomised to tailored internet based cognitive behaviour therapy, 11 (33) %) completed all prescribed modules within the treatment period, 18 (55 %) completed 75 per cent of the prescribed modules and 22 (67 %) completed 50 per cent of the prescribed modules. A subsample of the 33 participants allocated to the treatment group (n = 19) completed a 1-year follow-up. On the primary outcome measure BAI, a score of 10.37 (SD = 6.83) was obtained, suggesting that improvements were sustained (within-group Cohen's d = 0.80). Similar effects were found for the GAD-7, PHQ-9, MADRS-S, CORE-OM and OOLI, with within-group effect sizes ranging from d = 0.63 to 1.13. There were significant partial correlations between pre-treatment scores on the WCST perseverative errors and change scores on the main outcome BAI (r = -.45, p = .05, n = 20), and on CORE-OM, MADRS-S, QOLI, PHQ-9 and GAD-7. There were no significant partial correlations for the BAI (r = .20, p = .38, n = 22), CORE-OM, MADRS-S, OOLI, PHO-9 or GAD-7 when a similar analysis was conducted on the selfreported CFQ.

The results of this study are consistent with several clinical trials on internet-based cognitive behavior therapy for older adults (Dear et al., 2013; Dear et al., 2015a, b; Titov et al., 2015; Zou et al., 2012) and represents one potential approach for increasing access to treatment for the age group.

To date, while consistent results have been obtained across trials, all of the research on ICBT programs specifically designed for older adults has been conducted by a limited number of research groups and the clinical trials have involved relatively small numbers of participants. Alongside this, there is also a need for studies examining the implementation of ICBT for older adults into routine practice, particularly where participants are referred to ICBT treatment rather than seeking ICBT as is often the case in clinical trials.

General discussion

Main findings and conclusions

The aims of this thesis were to further develop and test the effects of tailored internet-based cognitive behavioural therapy on the basis of age, for adolescents, young adults and older adults. Specifically by developing and testing the effects of individually tailored internet-based cognitive behavioural therapy for adolescents with anxiety and comorbid depressive symptoms and by adapting and testing the effects of individually tailored internet-based cognitive behavioural therapy for young adults and older adults with anxiety and comorbid depressive symptoms. These aims were tested in two pilot effectiveness studies and two efficacy randomised controlled trials. In these trials modules from previous trials on tailored ICBT (Carlbring et al., 2011; Johansson et al., 2012; Bergman Nordgren et al., 2014) were adapted for the different age groups both in regards to language and with clinical examples, with different fictitious case examples but sharing the same interventions. The modules are all based on established and evidence-based CBT interventions and modules contain relevant components like psychoeducation, exposure exercises and behavioural experiments. The treatment can be considered empirically driven rather than theory driven; with modules origin from diagnose specific ICBT protocols on panic disorder (Carlbring et al., 2006), generalized anxiety disorder, social phobia (Andersson et al., 2006) and depression (Vernmark et al., 2010).

Based on the results from the first (Paper I) and second (Paper III) pilot effectiveness trial a tentative conclusion is that online materials based on CBT could be useful along with standard treatment delivered in child and adolescent psychiatric clinics and in youth care. These are the first ICBT studies of its kind using an adolescent and youth sample in a clinical setting in Sweden. The significant effects that were found on all dependent measures immediately following treatment and adherence rates for the adolescents are consistent with previous trials on CBT and ICBT for adolescents and slightly higher than adherence rates for young adults.

As was shown in Paper II, significant treatment effects were found for all dependent measures immediately following treatment, and significant time effects at the 12-month follow-up, showing that a majority of the participants remained stable after completing their treatment which indicates that tailoring an internet-based treatment can be a feasible approach in the treatment of panic symptoms and comorbid anxiety and depressive symptoms. The results indicate that younger adults benefit from treatment as much as adults between 30 and 45 years of age. The prevalence of untreated anxiety and depression represents a critical public health issue in the different age groups, and the proportion of older adults is set to grow over the coming decades. Adolescents, young adults and older adults have until recently been overlooked in global health and social policy, one reason why they have had fewer health gains with economic development than other age groups. Studies in adults suggest that most mental disorders begin before 25 years of age, most often between 11 and 18 years (Patton et al., 2016). Recent prospective studies have found that while mental health problems are very common during adolescence, not all persist into adulthood, particularly if the episodes are brief. These understandings have put a growing emphasis on early clinical interventions through more accessible and better resourced primary health care through youth focused mental health services (Halje et al., 2015). Online and mobile-phone interventions could play a positive part in prevention and promotion of access to clinical services since despite their clinical efficacy; there are numerous barriers to traditional face-to-face psychological treatments. Internet-based ICBT represents one potential approach for increasing access to treatment and several clinical trials have now been conducted with encouraging results (i.e. Hedman et al., 2012).

The findings of Paper IV, the first of its kind in Sweden, indicate that tailored ICBT is beneficial for older adults age 60 and above, and previous trials in Australia and Canada indicate that ICBT is acceptable to older adults and can result in clinically significant reductions in levels of anxiety and depression. In this paper, significant partial correlation between perseverative errors on the WCST and change scores on the included outcome measures was found. Further studies using WCST when testing

ICBT for older adults is needed to consolidate the predicting findings in this study that are preliminary and based on a small sample.

Limitations

There are limitations to this thesis. In regards to Paper I, the significant results and large within-group effect sizes reported in this study should be viewed with caution given the most important limitation of this study: the small sample size and the lack of randomization to a control group, limiting the internal validity of the study. In addition, there was no follow-up of treatment results after the post-assessment. However, the study was conducted in a clinical setting with a population that is common in child and adolescent psychiatry. Inclusion criteria were inclusive, with few restrictions with regards to comorbidity, thus strengthening the external validity. This means that the results are interesting from the standpoint of generalizability, and further studies ought to be conducted with larger samples. Tailored ICBT seems to be a way to reach adolescents who had not entered psychological treatment before and ICBT may be useful in reaching adolescents reluctant to attending the clinic for treatment. In Paper II the prescription of treatment modules in the study may be unreliable because it was based on a structured diagnostic procedure (SCID-I) and clinical impression by relatively inexperienced clinicians. A more comprehensive clinical assessment, such as a functional analysis, may have resulted in the prescription of other modules. Also worth mentioning is that the treatment modules consisted of modules derived from diagnosis-specific trials and were mainly structured after each diagnosis (i.e. panic disorder and not panic symptoms). This was addressed in the other trials (Paper I, II and IV), the modules were made symptom specific and less difficult, and less specific for particular diagnoses. Second, the use of a waitlist control condition is a limitation of Paper II. The use of a passive control group means that the effect of nonspecific factors cannot be determined. The lack of a comparison group at the 12-month follow-up makes it difficult to conclude that the improvements over the follow-up period were due only to the effects of the individually tailored treatment. Another limitation is the size of the study, which in turn affects the generalizability of the results. Another limitation is that the participants in this study had expressed an

interest in ICBT for their problems and therefore the participants may have been highly motivated to undergo treatment. However, we did not measure how highly motivated the participants were. A final limitation of this study was its inability to detect age differences, and it is possible that small but clinically meaningful differences would have been detected with a larger sample. The results of Paper III are in line with Halje et al. (2015) who examined CBT treatment at Youth Health Care Centers within the same county where this study took place. The results should be viewed with caution due to a small sample size and the lack of randomization, however this was a pilot effectiveness study with experienced psychologists conducting the interviews and administrating the treatment, which strengthens the validity. The use of standardized diagnostic interviews is not common practice at Youth Health Care Centers in the county (Halje et al., 2015). In Paper IV, the sample was small. There was not a pure waitlist control group and the control group was provided with support, with the role of support unclear. In the field of depression, supportive non-directive therapy has been found to have positive effects (Cuijpers et al., 2012). This may however also be regarded as a strength of the study as it is well established that psychological treatment, including ICBT, is better than just waiting (Cuijpers & Cristea, 2016). Second, the treatment approach involved selecting treatment modules based on symptoms and preferences and the reliability of the selection procedure is not established. Preliminary work implies that clients may be able to take more responsibility for the selection of treatment modules (Andersson et al., 2011), but it could also be the case that some components like exposure are more easily avoided. Overall it is a limitation that it is unclear which modules are driving effects, and also a problem with measures as transdiagnostic and tailored approaches would benefit from tailored assessments. The sample was recruited via mass media and even if this particular study had a lower proportion of participants with university education than we usually see (about half as compared to two thirds in many previous studies), it is still the case that we cannot infer that this treatment would work under regular clinical conditions. On the other hand ICBT for adults has been found to work in effectiveness studies (Andersson & Hedman, 2013), and in the study by Mewton et al. (2013) this appeared to be the case for older adults as well. One limitation that all the trials have are the measurements used to measure

anxiety and depression. The measurements are general measures of anxiety and does not target specific anxiety disorders. The measurements are also not adapted for the different age groups in general with the exception of CATS, although some are validated for the age group. In the future it is recommended to include age and diagnostic specific measurements.

Future Directions

Despite the positive findings of the studies in this thesis, there is a need for more research examining the acceptability and effectiveness of ICBT for adolescents, young adults and older adults with anxiety and depression. While consistent results have been obtained across these trials, all of the research on tailored ICBT has been conducted by one research group and the trials have involved relatively small numbers of participants. There is, therefore, a strong need for independent replication by other research groups, and the implementation and evaluation of larger trials involving much larger numbers of participants to establish the robustness of observed clinical effects. Alongside this, there is also a need for studies examining the implementation of tailored ICBT for different age groups into routine practice, particularly where participants are referred to ICBT treatment rather than seeking ICBT. It would be interesting to see if a briefer version of ICBT, including psychoeducational components, could increase motivation to seek face-to-face CBT and reduce risks of possible negative effects in digital distributed treatments (Rozental et al., 2014). Another suggestion would be to use an internet platform for homework assignments as part of regular CBT (Månsson et al., 2013), in other words, blending face-to-face and internet components. Moreover, given the smaller number of participants in existing trials, there has not been the ability to examine whether the various age cohorts of age groups differ in terms of how acceptable they find tailored ICBT or how they respond to tailored ICBT. Related to this issue, there are limited empirical data to clearly indicate whether, and if so when, different age groups might require or might benefit from ICBT programs designed specifically for them (i.e. providing ageappropriate examples), that is, compared with programs designed generally for adults with anxiety and depression. The degree to which internet-based treatments need to be modified for different age groups holds important

implications for the implementation of ICBT into routine care. Before the implementation on a larger scale of internet-based cognitive behavioural therapy in primary care and psychiatric settings for adolescents, young adults and older adults more effectiveness trials are highly recommended. The process of tailoring the treatment is also an area that can be examined further, perhaps sophisticated algorithms can be explored for the tailoring process. Results comparing tailored ICBT with transdiagnostic ICBT indicates that the treatments have similar effects (Păsărelu et al., 2016) and it would be interesting to directly compare the different treatment formats. Other aspects such as mechanisms of change and predictors for tailored ICBT need to be explored further.

Consequently, there is a significant opportunity and considerable benefit in future research exploring tailored ICBT for treating anxiety in different age groups across the life span.

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