

# Digital stress management in cancer: Testing StressProffen in a 12-month randomized controlled trial

Elin Børøsund, PhD <sup>1</sup>; Shawna L. Ehlers, PhD <sup>2</sup>; Matthew M. Clark, PhD <sup>2</sup>; Michael A. Andrykowski, PhD <sup>3</sup>; Milada Cvcancarova Småstuen, PhD <sup>4</sup>; and Lise Solberg Nes, PhD <sup>1,2,5</sup>

**BACKGROUND:** Cognitive-behavioral stress management interventions are associated with improved psychological well-being for cancer survivors. The availability of, access to, and outreach of these in-person interventions are limited, however. The current study, therefore, evaluated the efficacy of StressProffen, a digital application (app)-based stress management intervention for cancer survivors, in a 12-month randomized controlled trial. **METHODS:** Cancer survivors 1 year or less after their treatment (N = 172) were randomized to the StressProffen intervention (n = 84) or a usual-care control group (n = 88). The intervention was delivered in a simple blended care model: 1) 1 in-person introduction session, 2) 10 app-based cognitive-behavioral stress management modules, and 3) 2 follow-up phone calls. Stress (Perceived Stress Scale), anxiety and depression (Hospital Anxiety and Depression Scale), self-regulatory fatigue (Self-Regulatory Fatigue 18), and health-related quality of life (HRQOL; RAND-36) were examined at the baseline and at 6 and 12 months. Generalized linear models for repeated measures were fitted to compare effects over time. **RESULTS:** Participants were mainly female (82%), had a mean age of 52 years (standard deviation, 11.3 years; range, 20-78 years), and had a variety of cancer types (mostly breast cancer [48%]). Over the 12-month study time, the intervention group reported significantly decreased stress ( $P < .001$ ), depression ( $P = .003$ ), and self-regulatory fatigue ( $P = .002$ ) as well as improved HRQOL (for 6 of 8 domains,  $P \leq .015$ ) in comparison with controls. The largest favored effects for the intervention group were observed at 6 months: stress (estimated mean difference [MD],  $-5.1$ ;  $P < .001$ ), anxiety (MD,  $-1.4$ ;  $P = .015$ ), depression (MD,  $-2.1$ ;  $P < .001$ ), self-regulatory fatigue (MD,  $-4.9$ ;  $P < .001$ ), and HRQOL (7 of 8 domains;  $P \leq .037$ ). **CONCLUSIONS:** Digital stress management interventions such as StressProffen have the potential to extend the outreach of psychological interventions and provide easily available and effective psychosocial support for cancer survivors. *Cancer* 2021;0:1-10. © 2021 The Authors. *Cancer* published by Wiley Periodicals LLC on behalf of American Cancer Society This is an open access article under the terms of the Creative Commons Attribution-NonCommercial-NoDerivs License, which permits use and distribution in any medium, provided the original work is properly cited, the use is non-commercial and no modifications or adaptations are made.

**KEYWORDS:** cancer survivors, cognitive behavioral, electronic health (eHealth), mobile applications, psychological distress, psycho-oncology, stress management.

## INTRODUCTION

Cancer diagnosis and treatment are associated with physical and psychosocial challenges, including discomfort, fatigue, pain, stress, distress, worry, anxiety, and depression.<sup>1-3</sup> Quality of life (QOL)<sup>1-6</sup> and the capacity to control or alter (ie, self-regulate) thoughts, feelings, and behavior<sup>7,8</sup> are often also affected, and coping during and after cancer can be challenging.<sup>3,5,8,9</sup>

### *Psychosocial Interventions for Stress Management and Coping in Cancer*

Psychosocial interventions aiming to support stress management and coping have for decades been shown to promote well-being, including improved QOL and reduced stress, distress, anxiety, and depression for cancer survivors.<sup>2,4,5,10-15</sup> There are also some indications that the positive impact of such interventions may last for more than 10 years.<sup>16</sup>

Access barriers to in-person psychological interventions exist, however; they include availability, the geographical distance from the intervention site, costs/limited insurance coverage, and cancer survivors not feeling well enough or comfortable enough to participate in in-person sessions.<sup>5,15</sup> In light of continued findings that cancer survivors have many unmet needs, including needs for psychosocial support and care,<sup>5,17</sup> health care delivery methods with higher accessibility and outreach are needed.

**Corresponding Author:** Elin Børøsund, PhD, Department of Digital Health Research, Division of Medicine, Oslo University Hospital, Pb 4950 Nydalen, 0424 Oslo, Norway (elin.borosund@rr-research.no).

<sup>1</sup>Department of Digital Health Research, Division of Medicine, Oslo University Hospital, Oslo, Norway; <sup>2</sup>Department of Psychiatry and Psychology, College of Medicine and Science, Mayo Clinic, Rochester, Minnesota; <sup>3</sup>Department of Behavioral Science, College of Medicine, University of Kentucky, Lexington, Kentucky; <sup>4</sup>Faculty of Health Sciences, Oslo Metropolitan University, Oslo, Norway; <sup>5</sup>Institute of Clinical Medicine, Faculty of Medicine, University of Oslo, Oslo, Norway

We thank the cancer survivors participating in this study, the participating health care providers at Oslo University Hospital who aided with study recruitment, and the project study support team at the Department of Digital Health Research.

**DOI:** 10.1002/cnrc.34046, **Received:** July 9, 2021; **Revised:** October 1, 2021; **Accepted:** November 13, 2021, **Published online** Month 00, 2021 in Wiley Online Library (wileyonlinelibrary.com)

### **Digital Psychosocial Interventions for Stress Management in Cancer**

Digital solutions in the form of electronic health (eHealth) interventions have the potential to enhance the delivery of health care and psychosocial support to cancer survivors. However, published results from randomized controlled trials (RCTs) testing psychosocial eHealth interventions for cancer survivors are mixed and inconclusive.<sup>18-22</sup> Examining these findings, researchers have pointed to a need for eHealth interventions for cancer survivors to be theoretically based; to have significant involvement from end users (ie, cancer survivors) and other key stakeholders (eg, health care providers) during the design and development phases; and to include high-quality outcome assessments, larger sample sizes, and longer follow-up periods to establish efficacy.<sup>19,20,22</sup>

### **StressProffen: A Digital Stress Management Intervention for Cancer Survivors**

In response to identified challenges with eHealth interventions in cancer, the current research team designed and developed StressProffen, an application (app)-based stress management intervention program in support of cancer survivors.<sup>23,24</sup> StressProffen combines elements from well-known cognitive-behavioral stress management strategies<sup>4,12,13</sup> and was designed and developed according to user-centered design methods with close collaborations between researchers, cancer survivors, health care providers (eg, psychosocial oncologists), and eHealth experts.<sup>23</sup> In line with recommendations to certify the feasibility of complex medical interventions,<sup>25</sup> a feasibility pilot study testing StressProffen revealed positive acceptability, usability, and feasibility, with positive indications related to decreased stress, anxiety, and self-regulatory fatigue as well as improved health-related quality of life (HRQOL) in cancer survivors.<sup>24</sup>

The initial RCT evaluation of the program (ie, at 3 months) showed significant pre-post between-group differences, with cancer survivors who received StressProffen reporting decreased stress and improved HRQOL in comparison with usual-care controls.<sup>26</sup> There was also a decrease in anxiety and depression in favor of the intervention group, although this was not statistically significant. Effect sizes in the 3-month evaluation were generally small, however, with large data variability, which may have contributed to this.<sup>26</sup> Also, although a 58% completion rate (ie, at least 7 of 10 modules<sup>26</sup>) is considered above average for eHealth interventions,<sup>27</sup> it is possible that 3 months was not enough time for the participating cancer survivors to

complete the intervention. The current study examined longer term results from the RCT and evaluated the efficacy of the StressProffen intervention program over a 12-month period. It was hypothesized that participants receiving the StressProffen intervention, compared with participants in a usual-care control group, would 6 and 12 months after the intervention initiation experience decreased perceived stress (the primary outcome) and decreased anxiety, depression, and self-regulatory fatigue and improved HRQOL (all secondary outcomes).

## **MATERIALS AND METHODS**

### **Design and Participants**

In this RCT, participants (ie, cancer survivors) were assigned to either 1) the app-based StressProffen stress management intervention or 2) a usual-care control group and were followed for 12 months.

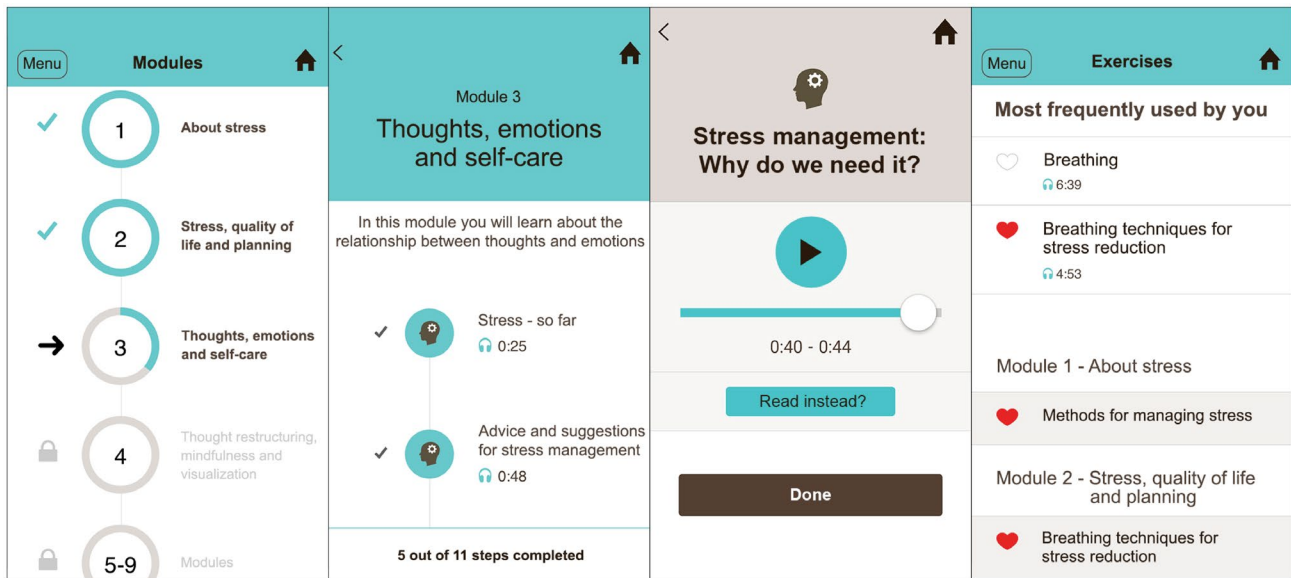
Patients diagnosed with any type or stage of cancer were recruited at a major medical center in Northern Europe or through social media from June 2017 to July 2019. The eligibility criteria were 1) currently or recently receiving cancer treatment (ie,  $\leq 1$  year after the completion of hospital treatment); 2) being 18 years old or older; 3) being able to speak, read, and understand Norwegian; 4) having access to a smartphone or tablet; and 5) being able and willing to attend 1 in-person introduction session. See also Børøsund and colleagues<sup>26</sup> for additional details on the study methodology.

The study was approved by the Regional Committee for Medical and Health Research Ethics (2016/14369) and the Hospital Privacy Protection Committee (2015/10204) and was registered with ClinicalTrials.gov (NCT02939612).

### **Study Procedure**

Patients were informed about the study via medical center social media and/or outpatient clinics/radiotherapy units, and they could request or self-initiate contact with the study team if interested. Participants provided written informed consent and completed baseline study questionnaires/outcome measures before randomization. Computerized randomization allocated study arms on a 1:1 basis (block size, 10) with stratification by gender and diagnosis (ie, breast cancer vs all other cancer diagnoses on the basis of pilot-study findings).<sup>24</sup>

All outcome measures and program use data were collected electronically through a secure server using an encrypted connection. Participants completed outcome



**Figure 1.** Examples of StressProffen screenshots.

measures at the baseline and at 3 (reported elsewhere),<sup>26</sup> 6, and 12 months.

### **The StressProffen Intervention**

In a simple blended care delivery model, the intervention group received the following: 1) an in-person introduction session with study personnel, 2) 10 app-based stress management modules (ie, StressProffen), and 3) follow-up phone calls 2 to 3 and 6 to 7 weeks after the introduction session. The introduction session was structured and served the dual purpose of introducing participants to the stress management concept<sup>1,10,12</sup> and helping participants to download and get started with the program.

The StressProffen program contains 10 modules with cognitive-behavioral and stress management educational material and exercises: 1) What is Stress; 2) Stress, QOL, and Planning; 3) Thoughts, Feelings, and Self-Care; 4) Mindfulness, Thought Challenging, and Guided Imagery; 5) Stress and Coping; 6) Social Support, Humor, and Meditation; 7) Anger Management and Conflict Style Awareness; 8) Assertiveness and Communication; 9) Health Behaviors and Setting Goals; and 10) Review and Summary.<sup>23</sup> See Figure 1 for examples of program screenshots. For more details about the content, development, and pilot testing of StressProffen, see Børø Sund and colleagues.<sup>23,24</sup>

Participants were encouraged to complete all 10 modules and to practice the content to become well acquainted with the material. The follow-up phone calls

were conducted by study personnel, structured, and included questions related to program impression and ease of use.

### **Data Collection and Outcome Measures**

A study-specific self-report questionnaire collected demographic, disease, and treatment information at the baseline. Comorbidity was measured at the baseline with the Self-Administered Comorbidity Questionnaire,<sup>28</sup> with total scores ranging from 0 to 57 and higher scores indicating a more severe comorbidity profile.

### **Psychosocial Outcome Measures**

#### **Primary outcome**

Perceived stress was measured by the Perceived Stress Scale, a 14-item scale measuring feelings and thoughts over the last month.<sup>29</sup> Items are rated on a 5-point Likert scale ranging from “never” (0) to “very often” (4). Total scores range from 0 to 56, with higher scores indicating higher perceived stress. Because it is not a diagnostic measure, the Perceived Stress Scale has no cutoff scores (ie, scores are labeled low, moderate, or high).

#### **Secondary outcomes**

Anxiety and depression were measured with the Hospital Anxiety and Depression Scale,<sup>30</sup> a 14-item measure of anxiety and depression, with 7 items measuring each subscale. Items are rated on a 4-point scale (0-3), with total scores ranging from 0 to 42 and with higher scores

indicating higher anxiety/depression. Score ranges are <8 (nonclinical), 8 to 11 (indicating the presence of anxiety/depression), and >11 (anxiety/depression). Nevertheless, there are some indications suggesting that these cutoff levels are too high for patients with cancer, and this may result in underrecognition of distress.<sup>31</sup>

Self-regulatory fatigue was measured with the Self-Regulatory Fatigue 18 (SRF-18), an 18-item self-report scale gauging the capacity to regulate cognitive, emotional, and behavioral components of self-regulation.<sup>32</sup> The SRF-18 contains 8 positively phrased items (eg, “I have no trouble making decisions”) and 10 negatively phrased items (eg, “I experience uncontrollable temper outbursts”). Items are scored on a 5-point Likert scale (1-5), with total scores ranging from 18 to 90 and with higher scores reflecting higher self-regulatory fatigue. The SRF-18 has acceptable internal consistency and reliability.<sup>32</sup>

HRQOL was measured with the 36-Item Short Form Health Survey (RAND-36 version),<sup>33,34</sup> a 36-item measure of physical, role, emotional, cognitive, and social function as well as physical health and general and global health/HRQOL. Subscale scores range from 0 to 100, with lower scores indicating higher disability (0 = maximum disability, 100 = no disability). A mean of 50 is generally considered normative for all subscales. The normative mean for the general Norwegian population may, however, be somewhat higher.<sup>35</sup>

### **Program Use**

Data related to program use (ie, app progress/activity) were extracted from user logs automatically collected and stored on a secure research server.

### **Statistical Analyses**

Baseline characteristics are summarized as means and standard deviations for normally distributed variables and as medians and ranges for variables with skewed distributions. Categorical data are presented as counts and percentages. For the analysis of between-group differences in outcome measurements, generalized linear models (GLMs) for repeated measures were fitted. To account for statistical dependencies as each individual was measured several times and time spans between the completed measurements varied, an unstructured covariance matrix was used to model variances. Models for each outcome consisted of 3 covariates: measurement (time), group, and interaction term (ie, time and group). All measured time points (ie, for outcome variables) were considered, and the GLM approach was, therefore, adjusted for baseline differences.

Because no statistically significant differences were observed between the intervention group and the usual-care control group in demographic- and disease-related variables at the baseline, no covariates were included in the GLMs. Outcome analyses were conducted with intention-to-treat analyses; all participants in each group were included independently of how much they had used the intervention. Between-group differences were computed as the intervention group change from the baseline to 6 and 12 months minus the control group change from the baseline.

Participants completing at least 70% of the modules (7 of 10) were defined as program completers.<sup>24,26</sup> Exploratory subanalyses for the intervention group only, using GLMs, were performed to detect potential differences in outcomes between intervention completers and noncompleters.

*P* values < .05 were considered statistically significant. Statistical analyses were completed with the Statistical Package for the Social Sciences (release 27; SPSS, Inc, Chicago, Illinois) and Stata (version 16).

## **RESULTS**

### **Sample Description**

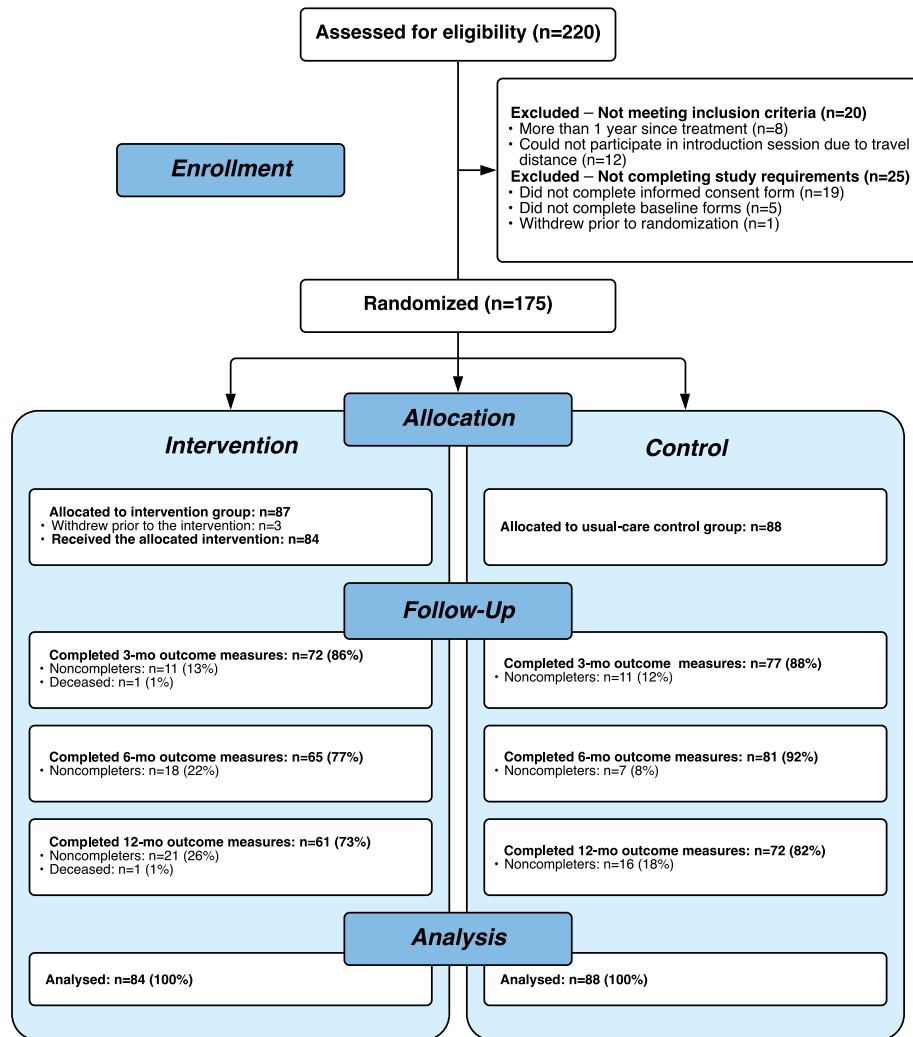
Among the 175 randomized participants, 3 intervention group assignees were unable to attend the in-person introduction session because of disease progression. The size of the final study sample was, therefore, 172. See Figure 2 for recruitment and retention details. Most participants were recruited by medical center clinic staff (102 of 172; 59%); the remainder were recruited through social media.

The mean age at inclusion was 52 years (standard deviation, 11.3 years; range, 20-78 years). The most common cancer type reported was breast cancer (48%). Most participants were female (82%), were married or cohabitating (70%), reported having a university/college education (81%), and were currently receiving sick leave/disability benefits (70%). See Table 1 for details.

### **Between-Group Differences**

Including measurements from all time points in the model showed statistically significant reductions in perceived stress, depression, and self-regulatory fatigue and improvements in 6 of 8 HRQOL domains over the 12-month study period for the StressProffen intervention group in comparison with the usual-care control group. See Table 2 and Figure 3 for details.

For all outcome measures, the largest intervention effects in favor of the intervention group were observed



**Figure 2.** Participant trial flow chart.

at 6 months. Between-group changes from the baseline to 6 months were statistically significant and showed that the intervention group improved on perceived stress, anxiety, depression, self-regulatory fatigue, and 7 of 8 HRQOL domains in comparison with the usual-care control group. See Table 2 for details.

### Program Use

Of the 84 participants in the intervention group, 57 (68%) completed at least 7 of the 10 modules within the 12-month study period. Thirty-nine (46%) completed all 10 modules. There were no significant differences in outcomes between intervention completers (ie,  $\geq 7$  modules) and noncompleters (ie,  $\leq 6$  modules). Participants used the program a median of 17.5 times (range, 3–170 times), and the median time from first

use to last use was 137 days (range, 10–365 days). At 6 months, 40% (34 of 84) still used the program. This declined to 21% (18 of 84) at 9 months and 12% (10 of 84) at 11 months.

### DISCUSSION

The current study was aimed at evaluating the efficacy of StressProffen,<sup>23,24,26</sup> a digital cognitive-behavioral stress management intervention for cancer survivors. The findings demonstrate a significant positive impact for the intervention group compared with usual-care controls, with between-group differences assessed over 12 months showing significant reductions in perceived stress (ie, the primary outcome) and significant reductions in depression and self-regulatory fatigue and



**TABLE 1.** Baseline Self-Reported Sociodemographic and Disease-Related Measures (N = 172)

Characteristic	Intervention Group (n = 84)	Control Group (n = 88)	P
Age, mean (SD), y	51.7 (10.5)	52.3 (12.0)	.725
Gender, No. (%)			.956
Female	69 (82)	72 (82)	
Male	15 (18)	16 (18)	
Marital status, No. (%)			.387
Married/cohabitating	56 (67)	64 (73)	
Single/divorced	28 (33)	24 (27)	
Education, No. (%)			.943
Elementary/high school	17 (20)	16 (18)	
University/college for ≤4 y	29 (35)	31 (35)	
University/college for >4 y	38 (45)	41 (47)	
Household annual income, No. (%) <sup>a</sup>			.629
<€40,000	9 (11)	9 (10)	
>€40,000–€60,000	15 (18)	17 (19)	
>€60,000–€80,000	5 (6)	11 (13)	
>€80,000–€100,000	17 (20)	14 (16)	
>€100,000	38 (45)	37 (42)	
Employment status, No. (%) <sup>b</sup>			.334
Full-time/part-time work	18 (21)	18 (21)	
Sick leave/disability benefits	61 (73)	59 (67)	
Retired/other	5 (6)	11 (13)	
Treatment, No. (%) <sup>c</sup>			
Surgery	66 (79)	60 (68)	.124
Chemotherapy	46 (55)	56 (64)	.236
Hormone therapy	21 (25)	23 (26)	.864
Radiation	34 (41)	40 (46)	.519
Immune therapy	8 (10)	10 (11)	.694
Other	10 (12)	14 (16)	.449
Diagnosis, No. (%) <sup>c</sup>			
Breast cancer	39 (46)	44 (50)	.639
Brain cancer	9 (11)	4 (5)	.126
Prostate cancer	6 (7)	4 (5)	.467
Other	30 (36)	36 (41)	.484
Metastases, No. (%)	12 (14)	11 (13)	.731
Months since diagnosis, median (range)	7.0 (0.25–120)	8.5 (0.25–240)	.183
Comorbidity, median (range)	3.0 (0–20)	3.0 (0–17)	.467

Abbreviation: SD, standard deviation.

<sup>a</sup>€1 is approximately US \$1.2 or approximately 10 Norwegian kroner (spring 2021).<sup>b</sup>Percentages do not add up to 100 because of rounding.<sup>c</sup>Participants could have received several treatments.

significant improvements in HRQOL (ie, the secondary outcomes).

Psychosocial interventions can support stress management and coping for cancer survivors,<sup>2,4,5,10–16</sup> but access barriers to such in-person interventions exist.<sup>5,15</sup> Findings from the use of StressProffen in the current study, therefore, provide great promise for digital solutions as supplements or alternatives to in-person psychosocial health care for cancer survivors. Even though the findings were statistically significant over time, more research is needed to explore the potential reasons why the largest intervention effects appeared around the

6-month follow-up. One explanation could be that psychosocial interventions have the most impact at an early stage in the cancer survivorship journey when the content is still new. If this was the case, however, findings at 3 months<sup>26</sup> could have been expected to be the most significant. Because participants were “on their own” progressing in the app (ie, it was user-driven rather than driven by provider guidance), however, it could be that 3 months was not enough time and that the 6-month follow-up revealed the benefits of more thoroughly acquired knowledge, skills, and strategies. A somewhat larger attrition rate in the intervention group compared with the control group from 3 to 6 months could potentially also have played a role if, for example, mainly particularly interested participants remained in the intervention at 6 months. Another factor to consider is that although both groups appeared to improve somewhat from the baseline to 3 months, the intervention group continued to improve, whereas the usual-care control group appeared to experience worsening symptoms from 3 to 6 months. It is possible that cancer survivors in the intervention group did in fact experience a sort of buffering effect from the StressProffen intervention during what may have otherwise been a challenging period in the cancer trajectory for most participants.

The simple blended care delivery model (ie, 1 introduction session, 10 app-based modules, and 2 follow-up phone calls) was used not only to ensure assistance should participants encounter technical issues but also to provide a sense of support throughout the self-management process. Even this simple form of blended care may have had an impact because guided eHealth interventions have been suggested to have better effects than self-guided interventions.<sup>20</sup> There was no follow-up contact between the research team and the participants between 6 and 12 months; if the simple blended care delivery induced a sense of guidance, this could have contributed to the more significant findings at 6 months. Another reason could be that the impact decreased as time progressed, and this supports the general notion that most interventions show declining impact over time. It should, however, be noted that even though a clear decline in program use was seen from 6 to 12 months, the impact of the program (ie, acquired knowledge and skills) could still be of benefit independently of actual app use.

Effect sizes in the current study ranged from small (<0.2) to moderate (<0.5). The great variation in data, indicating that some participants may have benefited more from the intervention than others, may have contributed to the limited study effect sizes. However, small

**TABLE 2.** Effects of StressProffen at 6 and 12 Months: Estimated Means From Generalized Linear Models

	Intervention Group (n = 84)		Control Group (n = 88)		Between-Group Differences <sup>a</sup>		<i>P</i>	Effect Size $\beta$	Time-Trend <i>P</i> <sup>b</sup>
	M	95% CI	M	95% CI	MD	95% CI			
Perceived stress (PSS-14)									<.001
Baseline	26.7	25.0-28.5	25.6	23.8-27.3					
6 mo	21.1	19.2-23.0	25.0	23.2-26.8	-5.1	-7.5 to -2.7	<.001	0.32	
12 mo	21.5	19.6-23.5	22.7	20.8-24.5	-2.3	-4.8 to 0.1	.063	0.14	
Anxiety (HADS-A)									.111
Baseline	8.5	7.6-9.4	8.8	7.9-9.7					
6 mo	6.6	5.6-7.6	8.3	7.4-9.2	-1.4	-2.5 to -0.3	.015	0.19	
12 mo	6.3	5.3-7.3	7.5	6.6-8.5	-1.0	-2.1 to 0.2	.101	0.14	
Depression (HADS-D)									.003
Baseline	5.4	4.6-6.1	5.0	4.3-5.7					
6 mo	3.5	2.7-4.3	5.3	4.5-6.0	-2.1	-3.0 to -1.1	<.001	0.36	
12 mo	3.8	3.0-4.6	4.5	3.8-5.3	-1.1	-2.0 to -0.1	.033	0.19	
Self-regulatory fatigue (SRF-18)									.002
Baseline	52.0	49.7-54.2	50.5	48.3-52.2					
6 mo	47.0	44.6-49.3	50.4	48.2-52.6	-4.9	-7.2 to -2.5	<.001	0.31	
12 mo	47.8	45.4-50.2	48.4	46.1-50.6	-2.0	-4.5 to 0.4	.102	0.13	
HRQOL (RAND-36)									
Physical functioning									.011
Baseline	72.9	68.4-77.4	79.2	74.8-83.6					
6 mo	79.4	74.5-84.2	76.5	72.0-81.1	9.1	2.9 to 15.4	.004	0.22	
12 mo	76.2	71.2-81.2	79.2	74.6-84.0	3.2	-0.3 to 9.6	.335	0.06	
Role-physical									.001
Baseline	19.9	11.7-28.2	37.5	29.4-45.6					
6 mo	40.8	31.6-49.9	33.8	25.5-42.2	24.5	11.4 to 37.6	<.001	0.29	
12 mo	34.4	25.1-43.8	46.3	37.6-55.0	5.7	-7.8 to 19.2	.406	0.06	
Bodily pain									.004
Baseline	57.5	52.4-62.8	64.1	59.0-69.3					
6 mo	66.6	60.8-72.4	59.6	54.3-64.9	13.6	5.8 to 21.4	.001	0.27	
12 mo	59.2	53.3-65.1	64.4	58.9-69.9	1.4	-6.6 to 9.4	.727	0.03	
General health									.075
Baseline	50.2	45.1-55.2	55.6	50.6-60.5					
6 mo	52.0	46.6-57.3	52.1	47.0-57.1	5.3	-0.1 to 10.4	.053	0.16	
12 mo	49.5	44.1-54.9	53.9	48.8-59.0	1.0	-4.5 to 6.5	.719	0.02	
Vitality									.001
Baseline	38.9	34.2-43.6	46.5	41.9-51.0					
6 mo	50.4	45.3-55.4	46.9	42.2-52.6	11.1	5.1 to 17.1	<.001	0.28	
12 mo	47.4	42.2-52.5	51.5	46.6-56.3	3.5	-2.7 to 9.7	.268	0.09	
Social functioning									<.001
Baseline	51.8	46.5-57.1	63.1	57.9-68.2					
6 mo	68.9	63.1-74.6	63.0	57.7-68.3	17.1	9.5 to 24.7	<.001	0.34	
12 mo	69.9	64.0-75.8	68.0	62.5-73.5	13.2	5.3 to 21.0	.001	0.26	
Role-emotional									.070
Baseline	46.8	37.7-55.9	52.7	43.7-61.6					
6 mo	64.0	53.9-74.2	53.7	44.5-62.9	16.1	1.0 to 31.2	.037	0.16	
12 mo	66.4	56.0-76.8	58.3	48.6-67.9	14.0	-1.6 to 29.5	.079	0.14	
Mental health									.015
Baseline	65.3	61.6-69.0	66.5	62.9-70.1					
6 mo	74.3	70.3-78.3	66.7	63.0-70.4	8.8	3.9 to 13.6	<.001	0.28	
12 mo	74.9	70.8-78.9	70.0	66.2-73.8	6.1	1.0 to 11.1	.018	0.19	

Abbreviations: CI, confidence interval; HADS-A, Hospital Anxiety and Depression Scale-Anxiety; HADS-D, Hospital Anxiety and Depression Scale-Depression; HRQOL, health-related quality of life; M, estimated mean; MD, estimated mean difference; PSS-14, Perceived Stress Scale; SRF-18, Self-Regulatory Fatigue 18. Three-month findings have been reported elsewhere (see Børøsd et al<sup>26</sup>).

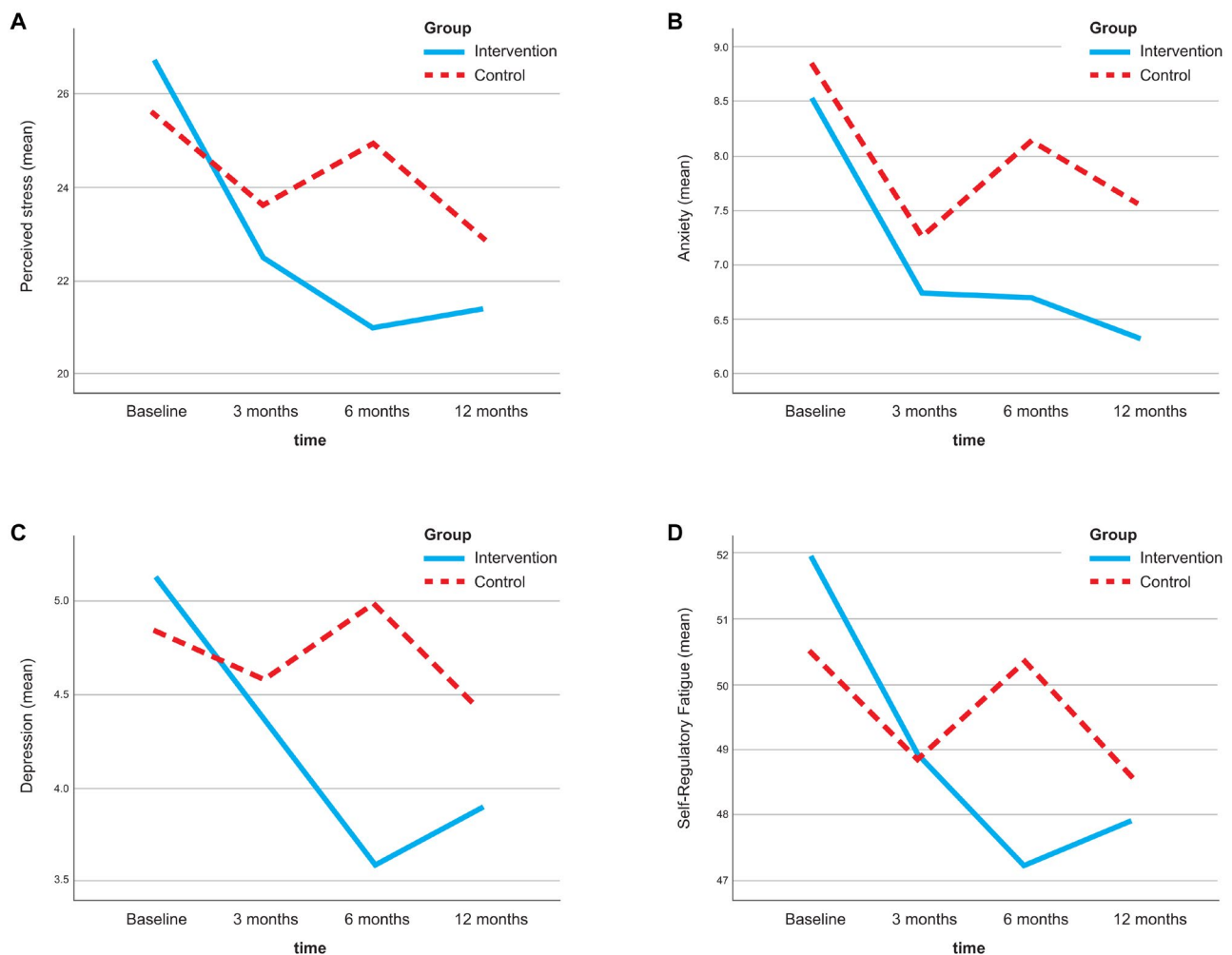
<sup>a</sup>Between-group differences were computed as the intervention group change from the baseline minus the control group change from the baseline. Negative values for PSS-14, HADS-A, HADS-D, and SRF-18 and positive values for RAND subscales indicate results in favor of the intervention group.

<sup>b</sup>Interactions between time and group (eg, a statistically significant *P* value indicates that the time trajectories were different for the 2 groups).

to moderate statistical effect sizes are not uncommon for psychosocial interventions in cancer and still provide evidence of statistical and clinical significance.<sup>11,14</sup> The fact that such effect sizes can be achieved with even minimal cost and effort via a digital approach further highlights

the significant potential for the outreach and impact of digital self-management interventions.

In addition to the longstanding limited availability of and access to psychosocial interventions for cancer survivors, a failure to meet the psychosocial needs of cancer



**Figure 3.** Estimated marginal means of (A) perceived stress (PSS-14), (B) anxiety (HADS-A), (C) depression (HADS-D), and (D) self-regulatory fatigue (SRF-18) for the intervention group ( $n = 84$ ) and the usual-care control group ( $n = 88$ ). Higher scores are indications of higher levels of stress, anxiety, depression, and self-regulatory fatigue. HADS-A indicates Hospital Anxiety and Depression Scale-Anxiety; HADS-D, Hospital Anxiety and Depression Scale-Depression; PSS-14, Perceived Stress Scale; SRF-18, Self-Regulatory Fatigue 18.

survivors during the recent coronavirus disease 2019 pandemic,<sup>17</sup> a pandemic bringing with it an instant need for new ways to deliver care, has also been indicated. Digital stress management solutions such as StressProffen may contribute to solving, or at least limiting, these types of challenges by providing innovative options for the outreach of effective psychosocial interventions even during challenging times.

To achieve the intended impact of interventions, adherence and continued use are vital. With adherence/completion rates for eHealth interventions sometimes as low as 20% to 40% (ie, 60%-80% attrition), however, the potential for intervention evaluation and effect is seriously compromised, and adherence and attrition surface as major

obstacles to the realization of eHealth interventions.<sup>21,27,36</sup> In the current study, as many as 68% of the participants could be considered completers (ie, completing  $\geq 7$  of 10 modules within the study period).<sup>24,26</sup> Completer status did, however, not affect outcomes, and this might indicate that even modest use of the StressProffen program could potentially have a positive impact.

The current study indicates that even minimal blended care models can strengthen delivery and the chance of impact. Future research should explore the extent needed to achieve such an impact. Could 1 phone call, or a simple introduction by a health care provider, be enough? Also, such explorations could determine whether completely user-driven digital self-management programs



can be effective; if so, this could mean even more simple, cost-effective health care. Future studies should also aim to rigorously compare well-established in-person psychosocial interventions for cancer survivors with comparable digital interventions such as StressProffen.

### Limitations

This study has several limitations. First, recruitment allowed for cancer survivors to contact the research team if they were interested in participation, and it can be assumed that these participants (41%) were particularly motivated; this potentially affected study generalizability. Second, disease status and treatment status were self-reported and could not be verified. Third, although a wide variety of cancer diagnoses were included in the current study, participants were mainly female, with a majority being breast cancer survivors. To strengthen generalizability and clinical utility, future research should aim to improve the gender balance and include more heterogeneous cancer survivor populations. Fourth, baseline scores were low to moderate for the majority of the outcome measures in the current study. It is possible that enrolling cancer survivors with higher baseline distress levels (ie, minimal distress score inclusion criteria) could have resulted in even higher intervention benefits for the participants. Finally, because of the access/not-access nature of the intervention, group allocation could not be blinded. Intervention group participants may, therefore, have anticipated potential effects, particularly if they were aware of findings from existing StressProffen publications.<sup>24,26</sup>

In conclusion, in an RCT, cancer survivors receiving StressProffen, a digital cognitive-behavioral stress management intervention delivered in a simple blended care model, compared with usual-care cancer survivor controls, reported improvements in perceived stress, depression, self-regulatory fatigue, and HRQOL over a 12-month time period. Digital stress management interventions such as StressProffen, built on evidence, with significant stakeholder involvement in the design and development process, have the potential to improve outreach and provide easily available and effective psychosocial support for cancer survivors. This type of care model could be especially effective in meeting distress management guidelines and accreditation standards in the provision of comprehensive cancer care.

### FUNDING SUPPORT

Funding was provided by the Norwegian Cancer Society (4602492-2013; principal investigator Lise Solberg Nes) and the Department of Digital Health Research of Oslo University Hospital (Oslo, Norway).

### CONFLICT OF INTEREST DISCLOSURES

Shawna L. Ehlers reports a role with Ask Mayo Expert. Lise Solberg Nes reports being an unpaid board member for dHealth AS. The other authors made no disclosures.

### AUTHOR CONTRIBUTIONS

**Elin Børø Sund:** Conceptualization, methodology, data collection, supervision, and writing (original draft, review, and editing). **Shawna L. Ehlers:** Conceptualization, methodology, and writing (review and editing). **Matthew M. Clark:** Conceptualization, methodology, and writing (review and editing). **Michael A. Andrykowski:** Conceptualization, methodology, and writing (review and editing). **Milada Cvancarova Småstuen:** Methodology and writing (review and editing). **Lise Solberg Nes:** Conceptualization, methodology, supervision, and writing (original draft, review, and editing). All authors gave final approval for the manuscript submission.

### REFERENCES

- Andersen BL, Kiecolt-Glaser JK, Glaser R. A biobehavioral model of cancer stress and disease course. *Am Psychol*. 1994;49:389-404.
- Stanton AL. Psychosocial concerns and interventions for cancer survivors. *J Clin Oncol*. 2006;24:5132-5137.
- Stanton AL. What happens now? Psychosocial care for cancer survivors after medical treatment completion. *J Clin Oncol*. 2012;30:1215-1220.
- Clark MM, Rummans TA, Atherton PJ, et al. Randomized controlled trial of maintaining quality of life during radiotherapy for advanced cancer. *Cancer*. 2013;119:880-887.
- Grassi L, Spiegel D, Riba M. Advancing psychosocial care in cancer patients. *F1000Res*. 2017;6:2083.
- Solberg Nes L, Liu H, Patten CA, et al. Physical activity level and quality of life in long term lung cancer survivors. *Lung Cancer*. 2012;77:611-616.
- Solberg Nes L, Ehlers SL, Patten CA, Gastineau DA. Self-regulatory fatigue in hematologic malignancies: impact on quality of life, coping, and adherence to medical recommendations. *Int J Behav Med*. 2013;20:13-21.
- Solberg Nes L, Ehlers SL, Patten CA, Gastineau DA. Self-regulatory fatigue, quality of life, health behaviors, and coping in patients with hematologic malignancies. *Ann Behav Med*. 2014;48:411-423.
- Stein KD, Syrjala KL, Andrykowski MA. Physical and psychological long-term and late effects of cancer. *Cancer*. 2008;112(suppl):2577-2592.
- Andersen BL, Farrar WB, Golden-Kreutz DM, et al. Psychological, behavioral, and immune changes after a psychological intervention: a clinical trial. *J Clin Oncol*. 2004;22:3570-3580.
- Andrykowski MA, Manne SL. Are psychological interventions effective and accepted by cancer patients? I. Standards and levels of evidence. *Ann Behav Med*. 2006;32:93-97.
- Antoni MH, Lehman JM, Kilbourn KM, et al. Cognitive-behavioral stress management intervention decreases the prevalence of depression and enhances benefit finding among women under treatment for early-stage breast cancer. *Health Psychol*. 2001;20:20-32.
- Gudenkauf LM, Ehlers SL. Psychosocial interventions in breast cancer survivorship care. *Breast*. 2018;38:1-6.
- Manne SL, Andrykowski MA. Are psychological interventions effective and accepted by cancer patients? II. Using empirically supported therapy guidelines to decide. *Ann Behav Med*. 2006;32:98-103.
- Rummans TA, Clark MM, Sloan JA, et al. Impacting quality of life for patients with advanced cancer with a structured multidisciplinary intervention: a randomized controlled trial. *J Clin Oncol*. 2006;24:635-642.
- Stagl JM, Bouchard LC, Lechner SC, et al. Long-term psychological benefits of cognitive-behavioral stress management for women with breast cancer: 11-year follow-up of a randomized controlled trial. *Cancer*. 2015;121:1873-1881.
- Archer S, Holch P, Armes J, et al. "No turning back" psycho-oncology in the time of COVID-19: insights from a survey of UK professionals. *Psychooncology*. 2020;29:1430-1435.

18. Buneviciene I, Mekary RA, Smith TR, Onnela JP, Bunevicius A. Can mHealth interventions improve quality of life of cancer patients? A systematic review and meta-analysis. *Crit Rev Oncol Hematol*. 2021;157:103123.
19. McAlpine H, Joubert L, Martin-Sanchez F, Merolli M, Drummond KJ. A systematic review of types and efficacy of online interventions for cancer patients. *Patient Educ Couns*. 2015;98:283-295.
20. Seiler A, Klaas V, Troster G, Fagundes CP. eHealth and mHealth interventions in the treatment of fatigued cancer survivors: a systematic review and meta-analysis. *Psychooncology*. 2017;26:1239-1253.
21. Chambers SK, Ritterband LM, Thorndike F, et al. Web-delivered cognitive behavioral therapy for distressed cancer patients: randomized controlled trial. *J Med Internet Res*. 2018;20:e42.
22. Penedo FJ, Oswald LB, Kronenfeld JP, Garcia SF, Cella D, Yanez B. The increasing value of eHealth in the delivery of patient-centred cancer care. *Lancet Oncol*. 2020;21:e240-e251.
23. Børøsund E, Mirkovic J, Clark MM, et al. A stress management app intervention for cancer survivors: design, development, and usability testing. *JMIR Form Res*. 2018;2:e19.
24. Børøsund E, Varsi C, Clark MM, et al. Pilot testing an app-based stress management intervention for cancer survivors. *Transl Behav Med*. 2020;10:770-780.
25. Craig P, Dieppe P, Macintyre S, Michie S, Nazareth I, Petticrew M. Developing and evaluating complex interventions: the new Medical Research Council guidance. *Br Med J*. 2008;337:a1655.
26. Børøsund E, Ehlers SL, Varsi C, et al. Results from a randomized controlled trial testing StressProffen; an application-based stress-management intervention for cancer survivors. *Cancer Med*. 2020;9:3775-3785.
27. Kelders SM, Kok RN, Ossebaard HC, Van Gemert-Pijnen JE. Persuasive system design does matter: a systematic review of adherence to web-based interventions. *J Med Internet Res*. 2012;14:e152.
28. Sangha O, Stucki G, Liang MH, Fossel AH, Katz JN. The Self-Administered Comorbidity Questionnaire: a new method to assess comorbidity for clinical and health services research. *Arthritis Rheum*. 2003;49:156-163.
29. Cohen S, Kamarck T, Mermelstein R. A global measure of perceived stress. *J Health Soc Behav*. 1983;24:385-396.
30. Zigmond AS, Snaith RP. The Hospital Anxiety and Depression Scale. *Acta Psychiatr Scand*. 1983;67:361-370.
31. Morse R, Kendell K, Barton S. Screening for depression in people with cancer: the accuracy of the Hospital Anxiety and Depression Scale. *Clin Eff Nurs*. 2005;9:188-196.
32. Nes LS, Ehlers SL, Whipple MO, Vincent A. Self-regulatory fatigue in chronic multisymptom illnesses: scale development, fatigue, and self-control. *J Pain Res*. 2013;6:181-188.
33. Ware JE Jr, Sherbourne CD. The MOS 36-Item Short-Form Health Survey (SF-36). I. Conceptual framework and item selection. *Med Care*. 1992;30:473-483.
34. Hays RD, Sherbourne CD, Mazel RM. The RAND 36-Item Health Survey 1.0. *Health Econ*. 1993;2:217-227.
35. Jacobsen EL, Bye A, Aass N, et al. Norwegian reference values for the Short-Form Health Survey 36: development over time. *Qual Life Res*. 2018;27:1201-1212.
36. Sieverink F, Kelders SM, van Gemert-Pijnen JE. Clarifying the concept of adherence to eHealth technology: systematic review on when usage becomes adherence. *J Med Internet Res*. 2017;19:e402.