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The Effect of Telehealth Interventions on Quality of Life of Cancer Patients: A Systematic Review and Meta-Analysis

Running Title: The Effect of Telehealth Interventions

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30 ABSTRACT

31 Introduction: In 2016, approximately 1.7 million new cases of cancer were diagnosed. Cancer
32 patients can have physical, functional, and psychosocial issues when dealing with cancer
33 treatment. Telehealth has been effectively introduced to help deliver treatment to patients
34 suffering from chronic disease, however, there is little consensus on its effectiveness in
35 administering socio-behavioral cancer treatments. Thus, this study determines the benefits of
36 telehealth-based interventions providing emotional and symptom support in improving quality of
37 life (QOL) among cancer patients.

38 Methods: Two researchers conducted comprehensive searches on PubMed, SCOPUS, Medline,
39 PsycINFO, ERIC, Psychology and Behavioral Collection, and Medline Complete. Key search
40 terms included telehealth or telemedicine and quality of life and cancer. Manuscripts were
41 included if they assessed a telehealth delivered intervention for adult cancer patients and
42 provided a QOL assessment. Data were extracted to calculate mean effect sizes for QOL
43 measures on the effectiveness of telehealth relative to usual care for cancer treatments.

44 Results: Out of 414 articles identified in our initial search, nine articles fit our inclusion criteria.
45 Both telehealth (Hedges $g = 0.211$, $p=0.016$) and standard of care (Hedges $g = 0.217$, $p<0.001$)
46 cancer treatment delivery methods demonstrated small but statistically significant improvements
47 in QOL measures. However, there were no statistically significant differences in effectiveness
48 between the telehealth interventions and usual care ($p=0.76$).

49 Conclusions: The results indicate that telehealth interventions are as effective at improving QOL
50 scores in patients undergoing cancer treatment as in-person usual care. Further studies should be
51 undertaken on different modalities of telehealth to determine its appropriate and effective use in
52 interventions to improve the quality of life for cancer patients undergoing treatment.

53

54

55 INTRODUCTION

56 Nearly two million new cases of cancer are diagnosed every year in the United States.¹ Cancer
57 patients can experience multiple issues during treatment, including physical, functional, and
58 psychosocial symptoms and complications.²⁻⁷ Cancer diagnoses can lead to severe psychological
59 distress and disrupt patients' lives, increasing strains on work, family, and social
60 relationships.^{2,8,9} Improved management of emotional distress and symptoms, especially after
61 new diagnoses and treatments, could significantly improve quality of life for cancer patients.¹⁰
62 Furthermore, the need for effective and cost-efficient interventions to address psychosocial
63 symptoms resulting from treatment will increase in the future with the aging demographic
64 distribution in the US and consequent increase in cancer diagnoses.¹¹

65
66 Telehealth has been effectively used to help manage many chronic conditions and to improve
67 compliance with treatment and patients' overall well-being.⁴ The terms 'telehealth' or
68 'telemedicine' are often used interchangeably and can have multiple definitions. Telemedicine is
69 often used to refer to diagnosis and monitoring technology, whereas telehealth may be used to
70 include management, education, and other allied health care services.¹² The Health Resources
71 and Services Administration defines telehealth as the use of technology to deliver health care,
72 health information, or health education at a distance.¹³ Telehealth technologies, including
73 telephone, videoconferencing, and internet-based interventions, have the capability of bringing
74 services into the patient's home and helping them manage their symptoms without needing to be
75 physically present at a hospital or clinic.^{3,10} Telemedicine patients have reported good acceptance
76 of and satisfaction with the use of technology in comparison with in-person visits.^{14,15} Providing
77 patients greater access to symptom management and emotional support services may lead to

78 patients taking a more active role in their health care and could improve patient outcomes
79 including overall quality of life (QOL).³

80

81 The purpose of this systematic review and meta-analysis is to examine the effect that telehealth
82 interventions providing emotional and symptom management have on cancer patients' QOL. To
83 our knowledge, there has been no study done to date that has examined the overall effect of
84 supporting patients in the management of their symptoms via telehealth technology in
85 comparison to in-person usual care (UC). We determine whether interventions utilizing
86 telehealth-delivered support are more effective in improving QOL versus UC from baseline until
87 the end of the intervention period.

88

89 METHODS

90 The recommendations outlined in the preferred reporting items for systematic reviews and meta-
91 analyses (PRISMA) statement were used to guide this systematic review and meta-analysis.¹⁶

92

93 2.1 Data Acquisition

94 An electronic database search was initially conducted from inception to December, 31 2016 by
95 two of the coauthors using the following databases: National Library of Medicine Catalog
96 (Medline/PubMed), SCOPUS, the Cumulative Index for Nursing and Allied Health Literature
97 (CINAHL), Ebsco Health (Medline complete). The initial key-term search consisted of:

98 “telehealth OR telemedicine” AND “Cancer” AND “quality of life OR assessment”. After the

99 initial search, article titles and abstracts were inspected for relevance to the inclusion and

100 exclusion criteria, followed by obtaining full-texts for identified manuscripts. Manuscripts were

101 then further scrutinized for inclusion and exclusion criteria post-retrieval. Reference lists of full-
102 text manuscripts were then hand-searched and cross-referenced for potentially relevant papers.
103 Another separate search on the Cochrane Library was conducted for systematic reviews
104 containing similar content. Relevant systematic reviews were then obtained and cross-referenced
105 for additional manuscripts missed during the original search. Consensus among all the authors
106 was then sought for an article's final inclusion in the meta-analysis.

107

108 2.2 Inclusion and Exclusion Criteria

109 All manuscripts included in the systematic review and meta-analysis must have been
110 published in a peer-reviewed journal and met the following inclusion criteria: 1) Patients
111 included must have had any form of cancer and been undergoing active treatment; 2) Patients
112 must have been adults, 18 years of age or older; 3) Interventions must have used some form of
113 telehealth/telemedicine, including but not limited to telephone calls and/or web based
114 interventions; 4) The focus of each intervention must have been on emotional support or self-
115 management of symptoms through counseling, educational intervention or telepsychiatry; and 5)
116 Studies must have used a measurable QOL scale or questionnaire. Studies were excluded if they:
117 1) Were written in a language other than English; 2) Included pediatric patients; 3) If they
118 assessed the efficacy of palliative care; or 4) Combined in-person and telehealth in the same
119 intervention.

120

121 2.3 Data extraction and analysis

122 For the systematic review portion of this study, descriptive data were extracted from each of the
123 included articles pertaining to their methodology and results. Numerical data extracted for the

124 meta-analysis included sample sizes, QOL measures means and standard deviations from
125 baseline and post-intervention as well as effect sizes for each study whenever data were
126 available. If effect size results were not reported, they were conservatively estimated based on
127 the obtainable data from each included study. Following the retrieval of study data, standardized
128 mean differences between baseline and post-test while adjusting for small sample bias (Hedges
129 g) were calculated for telehealth interventions and usual care (UC) separately.¹⁷ A mean effect
130 size (Δ) for both telemedicine and UC was determined using a random effects model due to the
131 uncertainty of evaluating a homogenous population.¹⁸ Heterogeneity was assessed via I^2 and Q -
132 statistics. To gauge the impact of bias from unpublished studies on the mean effect size, the fail-
133 safe N was also evaluated.¹⁸ All effect-size data and heterogeneity statistics were calculated with
134 the Comprehensive Meta-Analysis (V3.3.070, Biostat, Inc., Englewood, NJ) software package.
135 Effect size data were interpreted as 0.1-0.3=small, 0.3-0.5=moderate and >0.5 = large effects.¹⁹
136 After effect size calculations were acquired, independent t -tests were then used to determine if
137 differences existed between the effect sizes of the telemedicine and UC cancer delivery
138 interventions utilizing the IBM Statistical Package for the Social Sciences (SPSS) software
139 (Version 24.0, IBM, Inc., Armonk, NY). The significance level was set to $p < .05$ for all statistical
140 analyses a-priori.

141

142 RESULTS

143 Figure 1 is a flow diagram of our article selection process. Our initial search for articles using
144 our search terms within the designated literature databases yielded a total of 414 articles. A
145 search in the Cochrane database for systematic reviews containing similar content provided 5
146 systematic reviews, and all the references within the systematic reviews, totaling 370 article

147 titles, were screened. After titles and abstracts were screened, 57 articles were retained to be
148 assessed by two authors to ensure consensus on inclusion. After duplicates and those that did not
149 fit the inclusion criteria were excluded, full text assessments were performed on the 21
150 remaining articles. Nine articles were excluded due to the patient population being cancer
151 survivors and not in active treatment, and one article was excluded because the intervention was
152 exercise-based. Nine articles (Table 1) ultimately fit all systematic review and meta-analysis
153 criteria.

154
155 Five out of nine articles used telephone-based interventions (56%), another three studies used
156 web-based designs or connected devices (33%) and one (11%) utilized videoconferencing. The
157 time period for the studies varied, ranging from 6 weeks to one year. One article did not
158 specifically report the time period from baseline to final assessment, but stated it was one month
159 after treatment.²⁰ The mean age of the patients within the 9 articles ranged from 53 to 67 years of
160 age. Five of the articles focused on specific cancers, e.g., colorectal, breast, and head and neck
161 cancers,^{7,21–24} whereas three articles included three or more types of cancer within their study
162 population.^{20,25,26} Pfeifer et al. (2015) included both breast and prostate cancer patients.²⁷

163
164 Of the nine articles included, only one had statistically significant results for overall QOL scores
165 from baseline to end of the study period and did not have a comparable control group.²⁰ Two
166 articles did find clinically significant improvements in the intervention effect on QOL, but were
167 not statistically significant. Berry et al. (2014) did not find statistically significant between-group
168 changes in QOL overall, but did report statistical significance for a sub-analysis by age. There
169 was a statistically significant intervention effect for those ≥ 50 years of age, though not for those

170 younger.²⁶ Hegel et al (2011) found statistically significant improvements for the intervention
171 group compared to the usual care control for overall QOL as well as emotional and social well-
172 being subscales at 6 weeks. However, after the intervention was completed, the 12-week end of
173 study QOL scores were not statistically significant between groups.²² A similar effect was found
174 in Pfeifer's (2015) study, as there was no statistically significant difference between groups in
175 the overall QOL score, although there were statistically significant differences in physical well-
176 being after the intervention.²⁷ The Ruland (2013) and Rhyanen (2013) studies reported no
177 statistically significant between-group results for the telehealth intervention on QOL. Both
178 studies did find other statistically significant results related to lower anxiety and depression scale
179 scores for those in the intervention groups compared to the usual care controls over the study
180 period.^{23,24} Ruland et al. (2013) found that the intervention group had significant decreases in
181 depression scale scores and did not have the significant decreases that were found over time in
182 the control group for QOL and self-efficacy scores.²³ Rhyanen et al. (2013) collected data more
183 frequently and were able to associate QOL changes with events such as increases in QOL after
184 surgery and decreases in QOL at the end of radiotherapy.²⁴ The intervention group had a
185 continual decrease in anxiety over time, whereas the control group had greater anxiety before
186 surgery and chemotherapy, as well as during chemotherapy treatments.²⁴ In the study, anxiety
187 was statistically significantly associated with overall QOL scores and physical, psychological,
188 and spiritual well-being subscales.²⁴

189

190 In total, 16 individual effect sizes—nine for telehealth interventions and seven for UC—were
191 calculated. Across the nine studies included in the meta-analysis, 680 patients received
192 telehealth cancer interventions, while 602 patients received UC. The distribution for all

193 unweighted effect sizes calculated are displayed on a forest plot in Figure 2. The summary
194 statistics for the mean effect sizes for telehealth and UC with their 95% confidence intervals,
195 heterogeneity statistics and fail safe N calculations are reported in Table 2. Both telehealth ($\Delta =$
196 $0.211, p=0.016$) and UC ($\Delta = 0.217, p<0.001$) demonstrated small but statistically significant
197 mean effects compared to baseline QOL across the included studies. They each had relatively
198 low Q and I^2 values indicating homogeneity across the included studies (refer to supplemental
199 materials for funnel plot files). No statistically significant differences were present between the
200 mean effect sizes of telehealth and UC interventions ($t= -0.31, p=0.76$).

201

202 Multiple sensitivity secondary analyses were performed by revising the meta-analysis to include
203 only those articles that used the Functional Assessment for Cancer Therapy (FACT) scale for
204 quality of life. Including only these six studies increased the effect size of the intervention group
205 ($\Delta = 0.338, p=0.006$), and the effect size was considered moderate. The control group in this
206 analysis had a smaller increase in effect size ($\Delta = 0.256, p=0.013$). This suggests that using
207 different scales for measuring QOL may affect the measurable impact of the telehealth
208 interventions. We also stratified the meta-analysis to compare telephone interventions ($n=5$)
209 versus internet/device interventions ($n=4$). This resulted in telephone interventions having a
210 larger, moderate effect size ($\Delta=0.325, p=0.028$) than the internet/device interventions ($\Delta=0.092,$
211 $p=0.341$). However, these were not statistically different ($t= -0.584, p = 0.577$), likely due to the
212 low sample of studies.

213

214 DISCUSSION

215 Our study performed a systematic review of peer-reviewed studies that utilized telehealth
216 interventions to improve emotional support and symptom self-management for patients receiving
217 treatment for cancer. Our findings demonstrated a statistically significant, albeit small increase in
218 QOL for the telehealth intervention group relative to baseline across the nine studies in the meta-
219 analysis. The UC group had a similar, statistically significant improvement across seven studies,
220 but we found telehealth to be non-inferior to UC in improving quality of life for cancer patients.
221 Sensitivity analysis suggested that telephone-based interventions may be superior to
222 internet/device interventions for cancer patients.

223

224 The studies in our meta-analysis and systematic review were relatively homogenous as
225 demonstrated by funnel plots (refer to supplemental materials). Harrison et al. (2011) was the
226 only potential outlier showing significantly improved effectiveness of telehealth versus UC;
227 however, this was the only study analyzing patients with colorectal cancer.²¹ It is possible that
228 telehealth-based psychosocial treatments would vary in effectiveness across cancer diagnoses.
229 Unfortunately, there has been insufficient research to demonstrate this.²¹

230

231 Our findings are consistent with prior research demonstrating non-inferiority of tele-psychiatry
232 interventions versus face-to-face treatment.²⁸⁻³⁰ Thus, by maintaining a comparable QOL while
233 averting the need to travel for in-person therapy or treatment, the use of telehealth for
234 psychosocial support of cancer patients is likely to be cost-effective. Furthermore, telehealth may
235 be effective in improving outcomes other than QOL, however, such as patient satisfaction and
236 acceptability of the new modality.³¹ For example, a systematic review conducted by Calvin et al
237 suggested that most patients accept and are satisfied with many forms of telehealth interventions

238 they received.³² Another study showed that telehealth did not lead to lower patient satisfaction in
239 communicating with their providers.³³

240

241 Our inclusion criteria stated that all articles must have an overall quality of life measurement. Of
242 the nine studies in our systematic review, the majority (67%) used the FACT instrument—either
243 the general or cancer-specific FACT instrument—as shown in Table 1. The general FACT scale
244 was developed and validated between 1987 and 1992, in a five-phase process, including item
245 generation, item review and reduction, scale construction and piloting, initial evaluation, and
246 additional evaluation.³⁴ FACT-G is a 27-item instrument that has subscale scores for physical,
247 functional, social, emotional well-being and satisfaction with treatment.³⁴ Cancer-specific FACT
248 scales include those questions that are in the FACT-G but have additional questions that are
249 cancer specific, such as for colorectal cancer (FACT-C), breast cancer (FACT-B), and head and
250 neck cancer (FACT-HN).^{35–38} Berry et al.(2014) used the Symptom Distress Scale (SDS), which
251 has 15 items—the 13 included in the usual SDS instrument and an additional 2 questions related
252 to sexual activity and interest, and fever and chills.²⁶ The SDS used a 5-point Likert scale
253 ranging from no distress or normal (0) to severe distress (5), creating a total SDS score from an
254 unweighted summation of the scores.³⁹ Ruland et al. (2013) also used a 15-dimensional self-
255 administered instrument for measuring QOL based on similar symptoms to the SDS, but using a
256 5-point Likert score where higher scores denoted improved health status.^{23,40} The breast cancer
257 version of the Quality of Life Instrument was used by Ryhanen and colleagues.²⁴ Their
258 instrument had 46 items grouped into four subscales related to physical, psychological, social
259 and spiritual well-being. A 10-point Likert scale was used with 0 indicating the worst outcome

260 and 10 the best outcome. An overall QOL score was created by summing the subscale variables
261 and calculating the mean values.⁴¹

262

263 Although our focus was on the improvement in quality of life of cancer patients who received a
264 telehealth intervention for emotional or symptom management support, it is important to note
265 that telehealth increases access to care for cancer patients, as well as for those suffering from
266 other chronic conditions. Rural patients are at higher risk for decreased access to specialized
267 care, and telehealth has been found to increase access to quality care. Telehealth can ease the
268 burden of travel time, cost, and the discomfort that may be associated with long travel times.^{42,43}
269 Telehealth can also overcome issues related to ethnicity, culture, and language that affect health,
270 by facilitating access to culturally competent providers and interpreters.⁴³

271

272 Our study should be interpreted in the context of certain limitations. Our meta-analysis had a
273 small sample size of manuscripts and patient pools, and thus we were unable to perform a
274 moderator analysis to determine if alternative factors influenced the effectiveness of treatment
275 delivery. On a similar note, different cancers, stage of cancer and treatment protocols may have
276 varying impacts on QOL, which we were unable to explore due to the limited number of studies.

277 ⁴⁴⁻⁴⁶ A study on the factors affecting the quality of life of cancer patients undergoing
278 chemotherapy found worse quality of life in breast cancer, head and neck, sarcoma, lung and
279 gynecological cancers. Colorectal cancer patients were found to have the better quality of life.⁴⁶

280 Lower quality of life in breast cancer patients may be due to changes in self-image due to
281 surgery and hair loss, as well as decreased sexual function and early menopause. Similarly, head
282 and neck cancers and sarcomas surgical treatment can lead to disfigurement and cause lower

283 quality of life for patients.⁴⁶ In addition, we were unable to effectively assess manuscript quality
284 as part of our analysis as there was a range of study designs included. Due to the low sample, we
285 chose to include all studies relevant to our inclusion criteria and agreed upon by author
286 consensus, regardless of design. Because of inconsistent and limited published data, we erred on
287 the conservative side when necessary during effect size calculations; however, this only occurred
288 for two studies, Pfeifer et al. (2015) and Hegel et al. (2011).^{22,27}

289

290 CONCLUSIONS

291 Our systematic-review with meta-analysis demonstrated that supplementary interventions
292 through telehealth have a comparable impact on quality of life scores relative to in-person usual
293 care. Utilizing telehealth, may allow clinicians and healthcare systems to increase access for
294 those cancer patients who lack the means to travel for additional treatment or are rurally located
295 creating increase travel costs and time. Some of the studies in this meta-analysis did see
296 improvements in other areas such as depression, anxiety, and emotional, social, and physical
297 well-being, even when overall quality of life was not statistically significantly improved. Our
298 findings suggest more studies need to be conducted on the impact of telehealth interventions
299 across different cancer diagnoses to gain better insight into the differential effect these
300 interventions may have on quality of life for cancer patients undergoing treatment.

301

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303

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305

306

307 AUTHOR DISCLOSURE STATEMENT

308

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