Palliative and Supportive Care (MP Davis, Section Editor)



The Effect of Prognostic Communication on Patient Outcomes in Palliative Cancer Care: a Systematic Review

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Published online: 23 April 2020 © The Author(s) 2020

This article is part of the Topical collection on Palliative and Supportive Care

Keywords Advanced cancer · Palliative care · Prognosis · Truth disclosure · Physician-patient communication · Patient outcomes

Abstract

Background While prognostic information is considered important for treatment decision-making, physicians struggle to communicate prognosis to advanced cancer patients. This systematic review aimed to offer up-to-date, evidence-based guidance on prognostic communication in palliative oncology. Methods PubMed and PsycInfo were searched until September 2019 for literature on the

Methods PubMed and PsycInto were searched until September 2019 for literature on the association between prognostic disclosure (strategies) and patient outcomes in palliative cancer care, and its moderators. Methodological quality was reported.

Results Eighteen studies were included. Concerning prognostic disclosure, results revealed a positive association with patients' prognostic awareness. Findings showed no or positive associations between prognostic disclosure and the physician-patient relationship or the discussion of care preferences. Evidence for an association with the documentation of care preferences or physical outcomes was lacking. Findings on the emotional consequences of prognostic disclosure were multifaceted. Concerning disclosure strategies, affective communication seemingly reduced patients' physiological arousal and improved perceived physician's support. Affective and explicit communication showed no or beneficial effects on patients' psychological well-being and satisfaction. Communicating multiple survival scenarios improved prognostic understanding. Physicians displaying expertise, positivity and collaboration fostered hope. Evidence on demographic, clinical and personality factors moderating the effect of prognostic communication was weak.

Conclusion If preferred by patients, physicians could disclose prognosis using sensible strategies. The combination of explicit and affective communication, multiple survival scenarios and expert, positive, collaborative behaviour likely benefits most patients. Still, more evidence is needed, and tailoring communication to individual patients is warranted. *Implications* Future research should examine the effect of prognostic communication on psychological well-being over time and treatment decision-making, and focus on individualising care.

Introduction

Although antineoplastic treatment options have evolved in the past decades, cancer remains a leading cause of death globally [1-3]. To a certain degree, physicians are able to gain insight into advanced cancer patients' prognosis and disclose the lifelimiting nature of the disease. However, recent therapeutic developments and associated altered disease outcomes challenge oncologists' prognostic assessments [4•]. Moreover, estimating an individual's life expectancy is undeniably complex [40, 5]. Nevertheless, in an era of patient autonomy and shared decision-making, physicians' provision of prognostic information is considered important for patients to make informed treatment choices [6–9]. In addition, disclosure of prognosis might be necessary for them to prepare for the end of life [10]. A vast majority of patients with metastatic cancer wishes to be informed about the expected outcome of their illness [11, 12].

Yet, internationally, half of patients with advanced cancer are not aware of their prognosis [13–15]. Patients with incurable cancer often misunderstand the palliative intent of their treatment and overestimate their life expectancy compared with their oncologists [7, 13, 16–

18]. What is more, some studies indicate that these misconceptions grossly remain unchanged over time as death approaches [19, 20].

A lack of prognostic awareness could lead to decisions in disaccord with patients' actual values [17]. While accurate prognostic understanding is commonly associated with a preference for comfort care [17, 21], engagement in advance care planning [22, 23] and improved quality of life [24–28], misunderstanding of prognosis is associated with the administration of aggressive anticancer therapy [16, 17, 22, 29] and lifesustaining treatment [23, 25] at the end of life. Additionally, documentation of end-of-life preferences [22, 30] and usage of hospice services [30–33] can be complicated by prognostic unawareness.

Prognostic unawareness might result from the way physicians and patients communicate [34]. Oncologists are often reluctant to discuss prognosis [15, 34–37] and seem to worry about damaging the physician-patient relationship, patients' hope or psychological well-being [6, 38, 39]. When oncologists provide prognostic information, current literature suggests that they use imprecise qualitative terms (e.g. "months to years") instead of

quantitative point estimates (e.g. means or medians) or survival rates (e.g. percentages) [34]. Furthermore, challenged with the fine art of balancing hope and honesty, physicians regularly emphasise the presence of beneficial prognostic markers, the best case scenario and "years" instead of "months" [34, 38, 40-42].

Most past research has focused either on the prevalence and consequences of prognostic unawareness or on patients' information preferences, using observational or exploratory methods [43•]. However, information on how to best engage in prognostic discussions is limited. More specifically, few studies investigate the independent effect of prognostic communication on patient outcomes in palliative oncology, or factors that influence this relationship [34]. Hence, existing guidelines are mostly based on descriptive studies or expert consensus. Finally, there is no up-to-date synthesis of literature on this specific topic [44, 45].

Comprehensive information about the effect of different approaches to prognostic communication on advanced cancer patients is needed. This effort could assist oncologists in these challenging conversations, optimise the delivery of prognostic information and enhance patient outcomes. Integrating knowledge on moderating factors could help to tailor communication to individual patients, and ultimately formulate evidencebased advice for physicians' clinical practice. Therefore, this systematic review addresses the following research questions:

- I. What are the effects of prognostic disclosure on patient outcomes in palliative cancer care?
- II. What are the effects of different strategies to prognostic disclosure on patient outcomes in palliative cancer care?
- III. What patient and context characteristics moderate the effect of prognostic communication on patient outcomes in palliative cancer care?

Methods

The Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) statement was used as a guideline for this systematic review [46]. No protocol was registered.

Literature search

From July till September 2019, PubMed and PsycInfo databases were searched for studies published in English, using no restriction on publication year. The following search terms were used as main index terms or free-text words: "prognosis" and "communication" or "physician patient interaction" and "neoplasms" and "palliative care". Additionally, synonyms and closely related words were used (Appendix 1). One recently published, not yet indexed article was found through a PubMed article alert.

Eligibility criteria

Original quantitative studies describing the association between physician-patient communication about prognosis and patient outcomes in palliative cancer care were eligible. Palliative cancer care was defined as care for patients with incurable, metastatic cancer, including end-oflife care. Studies with samples partially matching the target population were only included if subgroup results were available. Communication about prognosis was defined as communicating the absence of cure, terminal nature of the disease and/or life expectancy. Papers addressing "bad news" without specifying its definition did not suffice. Communication could be real (e.g. self-reported by physicians or patients or observed in medical records, audio-recorded consultations or individually adapted consultations) or hypothetical (e.g. manipulated in video-recorded or written vignettes). Studies qualified if the independent effect of (strategies for) prognostic disclosure on (any type of) patient outcome(s) was examined. Hence, studies investigating more general interventions, such as advance care planning, early palliative care, decisions aids, question prompt lists or communication skills training, were excluded. Qualitative and non-empirical research, case reports and studies investigating minors or caregivers were also excluded.

Study selection

Duplicates were removed. Two authors (NV, MM) double screened 10% of the resulting records based on title and abstract (N = 377). These authors independently agreed on the inclusion, exclusion or the necessity to retrieve full-text papers for 96% of this sample. For 4% (N = 15), judgement differed, but only on the necessity for further evaluation. The last author (IH) was involved to jointly decide on inclusion of papers with questionable eligibility and to specify the criteria. After resolving all differences, the remaining 90% (N = 3393) of records were screened individually based on title and abstract (NV, MM). Discussion between NV, MM and IH took place in case of doubt. These authors jointly evaluated potentially relevant papers in full text (N = 54) and decided on the final inclusion with 100% agreement.

Data extraction

Data were extracted independently by NV and MM with a standardised extraction form including first author, year of publication, country, study aims, design, sample (description and size), setting, type of prognostic communication (prognostic disclosure and/or disclosure strategy), definition of prognosis (predictor), assessment of predictor, patient outcome and assessment and relevant main and moderating effects (direction and significance with *p* values or confidence intervals). Interaction terms and predictors of patients' reactions to manipulated prognostic messages were regarded as moderating factors. The congruence of the independently extracted data was judged by MM. In case of doubt, discussion with NV, MM and IH took place until agreement was reached.

Quality assessment

Appendix 2 displays the quality assessment tools used. The adapted New-castle-Ottawa Scale was adjusted to assess the quality of non-experimental studies [47, 48]. Eight items were scored with 0 to 2 points, leading to a maximum score of 16. A quality checklist for experimental studies was self-developed to suit the various designs of the included studies. Items

were based on the adapted Newcastle-Ottawa Scale [47, 48], the Cochrane Consumer and Communication Review Group criteria [49, 50] and the Cochrane Collaboration's tool for assessing risk of bias [49]. Experiments using a within-subjects design were rated with 8 items (part A). Experiments using a between-subjects design were additionally assessed with 5 more items (part B). Items were scored with 0 or 1 point, leading to a maximum score of 8 or 13 points, respectively. Selfconstructed items assessing the clarity of the definition of prognostic communication were included in both quality assessment tools.

Total scores and percentages of the maximum score were calculated. Papers attaining < 50% of the maximum score were considered of relatively low quality. Scores of ≥50% reflected satisfactory quality. Importantly, mutual comparison is only appropriate for studies using similar designs.

Methodological quality was assessed by NV, MM and IH. After these authors reached consensus during double assessment of one third of the papers (N = 6), MM and IH evaluated the remaining papers independently (N = 12). In case of doubt, discussion between all assessors took place. Agreement on quality was reached for all studies.

Results

The search yielded 3770 non-duplicate records (Fig. 1). Eighteen papers were included. Study characteristics are shown in Table 1.

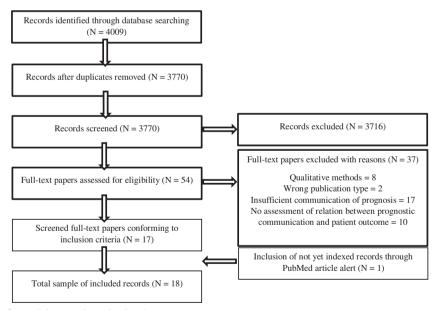


Fig. 1. Flow diagram for article search and selection

Authors (year, country)	Study aims	Design	Sample	Setting	Type of prognostic communication	Definition of prognosis (predictor)
Y. Aoki et al. (1997, Japan)	To examine how discourse of diagnosis, pathology and diagnosis, affects patients prognosis affects patients self-determination and attitude during the terminal stans.	Cross-sectional study	12 patients with metastatic lung or breast cancer or sarcoma	1 academic medical centre	Prognostic disclosure	Prognosis ¹ 7
E.H. Bradley et al. (2001, USA)	susges To examine (1) the proportion of advanced cancer patients who have a documented prognostic discussion in their medical records, (2) the potential factors associated with such discussions, (3) the nature of such discussions and (4) the association between such discussions and advance care alamina	Cross-sectional study	232 patients (2 65 years) with advanced brain, pancreas, liver, gall bladder or lung cancer	6 community hospitals	Prognostic disclosure	The word "prognosis", time frames until death, life expectancy and expected disease outcome
L.D. Gripe et al. (2012, USA)	To caracterize the parameter of a caracterize to a catual survival, patients' or oncologist's perceptions of prognosis or extent of prognosis or extent of prognosis or extent of prognosis or extent of prognosis caracterize concologist-reported extents or oncologist-reported extents or prognosis discussion moderate the association between patients' perceptions of prognosis and anxiety, or deassociation and anxiety or deassociation and anxiety or deassociation and anxiety.	Gross-sectional study	86 men with advanced cancer	1 cancer centre	Prognostic disclosure	Life expectancy
0.P., Danzi et al. (2018, Italy)	and analysely or legical solutions of supportive comments during a bad news consultation has a buffering effect on heart rate variability and (2) if it improves recall of provided information.	Experiment (between-subjects design, RCT)	60 healthy women without previous cancer history	Experimental setting	Prognostic disclosure and disclosure strategy: affective vs. standard communication	Incurability, life expectancy and treatment options
A.C. Finzinger et al. (2015, USA)	To examine (1) the proportion of patients who want to know their life expectancy and who report that their physician disclosed a prognostic estimate and (2) whether prognostic disclosure is associated with more accurate patient perceptions of life expectancy and more frequent advance care planning without harm to patients' well-being or the patient-physician relationship.	Cohort study	590 patients with metastatic solid metastatic solid manginancies with progressive disease after 2 1 line of palliative chemotherapy	(outpatient dinic)	Prognostic disclosure	Life expectancy
A.S. Epstein et al. (2016, USA)	To examine the effects of recent and past clinical prognostic discussions on changes in illness understanding	Cohort study	178 patients with advanced cancer refractory to prior chemotherapy whom oncologists expected to die within 6 months	9 cancer centres	Prognostic disclosure	Prognosis¹ and life expectancy

Table 1. (Continued)	ned)					
Authors (year,	Study aims	Design	Sample	Setting	Type of prognostic	Definition of prognosis
(former)	disconcordant views of prognosis and (2) the communication factors that may influence concordance about chance of cure		the patient was admitted to the intensive care unit or died within 12 months		statements about the past, present or future vs. uncertain statements	(manaid)
T. Rumpold et al. (2015, Austria)	To examine (1) the information preference of advanced lung cancer patients regarding cure rates and prognosis, (2) patients' satisfaction with an individually adapted medical consultation and (3) patients' emotional responses to the information	Quasi-experiment (allocation based on patient preference)	50 patients with advanced lung cancer	1 academic medical centre	Prognostic disclosure and disclosure strategy; qualitative vs. qualitative and quantitative information	Incurability and life expectancy
M.S.C. Sep et al. (2014, The Netherlands)	To ensurance the community of the commun	Experiment (between-subjects design, RCT)	50 healthy women without previous cancer history	Experimental setting	Prognostic disclosure and disclosure strategy: affective vs., standard communication	Incurability, life expectancy and treatment options
J.A. Sinn et al. (2016, USA)	To commitment definition of the depression, anxiety and perceptions of prognosis in patients with metastatic breast cancer and (2) whether symptom burden and prognostic understanding differed between patients receiving endocrine through and productine through and channothers and perceiving endocrine through and channothers are serving endocrine through an experience of the serving and channot are servin	Cross-sectional study	140 patients with metastatic breast cancer receiving either endocrine theapy (40) or chemotherapy (100)	1 cancer centre (outpatient clinic)	Prognostic disclosure	Prognosis ¹
L.M. Van Vliet et al. (2013, The Netherlands)	To examine the effect of more vs. less explicit prognostic information and reassurance about non-abandonment at the transition to paliative care	Experiment (within-subjects design)	104 patients with or survivors of breast cancer (51) and healthy women (53)	Experimental setting	Disclosure strategy: more vs. less explicitness and affective vs. standard communication	Life expectancy and treatment options
G.J. Wagner et al. (2010, USA)	To examine how provider communication and patient understanding of life-limiting illness relates to patient discussion of care preferences with providers and family by studying how often these elements of communication take place and studying the associations among them	Cross-sectional study (baseline of RCT)	400 inpatient veterans with a life-limiting illuess (26 having cancer, 224 having a non-cancerous disease)	1 veterans hospital	Prognostic disclosure	Life-limiting nature of the disease

	Relevant moderating effects³		NA A	₹
	Relevant main effects ²	or the choice of do-not-resuscitate order between prognostic disclosure vs. non-disclosure (p values NA) in hivarian analyces	Positive associations between prognostic discussion and do-not-resuscitate orders (95% CI = 1.1 to 4.2) and discussions about life-sustaining treatments (95% CI = 2.8 to 12.0) No association between prognostic discussion and advance directives (95% CI = 0.4	No association between patient-rated extent of prognostic discussion and anxiety (p = .318). In prognostic discussion and anxiety (p = .318). In bivariate analyses Negative association between patient-rated extent of prognostic discussion and depression (none vs. full discussions, p = .028); in multivariate analyses Interaction effect of patient-rated extent of prognostic discussions, p = .020); in multivariate analyses Interaction effect of patient-rated extent of prognostic discussion x patient-perceived prognosis on depression (more depression flowing a worse perceived prognosis with no or brief vs. full discussion, p = .011 vs. p = .188) Interaction effect of patient-perceived prognosis with full vs. no p = .002 vs. p = .444) No interaction effect of on anxiety (more anxiety following a worse perceived prognosis with full vs. no or brief discussion, p = .002 vs. p = .444) No interaction effect of on anxiety (more anxiety following a worse perceived prognosis with full vs. no or brief discussion, x patient-perceived prognosis on depression, or depression, or depression, or depression, or patient-rated extent of prognostic discussion x patient-perceived prognosis on depression, or depression, or patient-rated extent of prognostic
	Patient outcome and assessment	do-not-resuscitate order (registered in medical records)	Presence of do-not-resuscitate orders, discussions of life-sustaining treatment preference and advance directives.* (registered in medical records)	Anxiety and depression (HADS); patient perception of life sepectano; (survey question based on Weeks et al. [17, 18])
per)	Assessment of predictor	(registered in medical records)	Researcher-rated presence of prognostic discussion (registered in medical records)	Patient- and oncologist-rated extent of prognostic discussion (self-constructed survey question)
Table 1. (Continued)	Authors (year, country)	;	E.H. Bradley et al. (2001, USA)	L.D. Gripe et al. (2012, USA)

	Relevant moderating effects³	₹	NA
	Relevant main effects²	discussion × patient-perceived prognosis on anxiety (V aulers NA) High-frequency power decreased (P < .005), and low-frequency/ high-frequency and ingh-frequency and readment options, during accommunication about life expectancy and treatment options, there were higher rates of high-frequency power (P = .027) in the standard vs. affective condition in bivariate analyses, as well as more perceived doctor empathy (P = .037) and doctor support (P < .001) in the affective condition of the perceived recall ability or recall (p < .051), personal involvement, self-perceived recall ability or recall (p values NA) between the affective vs. standard condition after the manipulation in bivariate analyses. Interaction effect of the manipulation of the inter-bat interval with standard vs. A standard condition after the manipulation of the inter-bat interval with standard vs. affective communication, p = .002 and p = .030; better recall of additional treatment information following increased standard deviation frequency power with affective vs. standard communication, p = .002 and p = .030; better recall of information following increased high-frequency power with affective vs. standard communication, p = .007.	No difference in survival between prognostic disclosure vs. non-disclosure (p = .07) in bivariate analyses Positive association between prognostic disclosure and accuracy of life expectancy self-estimates within 3 months, 6 months and 12 months of actual survival (95% LI = 0.78 to 4.12: 95% LI = 1.11 to 3.73: 95% LI = 1.11 to 3.73: 95% LI = 1.11 to 3.51) in bivariate analyses, particulady preventing gross overestimation of > 2 years and > 5 years
	Patient outcome and assessment	Heart rate variability categorised in frequency and time parameters ⁸ (ECG signals from the ECG100C Electrocardiogram Amplifier); doctor empathy, doctor support, doctor engagement, personal involvement, self-perceived recal ability (self-constructed survey questions); recall (survey questions); recall (survey questions based on Sep et al. [53])	Survival (registered in medical records); prognostic understanding of life expectancy and health status (self-constructed interview questions); sad/depressed mood and woried/annious mood (MQOL psychological subscale); major depressive disorder and generalised anwiety disorder (SCID-5 and Endicott scale); patient-physician relationship (self-constructed interview questions)
nued)	Assessment of predictor	Providing reassurance or not during a simulated bad news consultation (video-recorded vignettess?)	Patient-rated presence of prognostic disclosure (self-constructed interview question)
Table 1. (Continued)	Authors (year, country)	(2018, Italy)	A.C. Enzinger et al. (2015, USA)

Patient outcome and assessment Patient outcome and assessment	(95% CI = 0.14 to 0.82; 95% CI = 0.08 to 0.47) Negative association between prognostic disclosure and length of life expectancy self-estimates (p = .0024) and positive association between prognostic disclosure and terminal illness and terminal illness acknowledgement (95% CI = 2.00 to 5.15) in multivariate analyses No associations between prognostic disclosure and sachess/depressed mood (p = .299), worning/disclosure and sachess/depressed mood (p = .259), worning/disclosure disclosure (95% CI = 0.83 to 4.14), generalised anxiety disorder (95% CI = 0.75 to 2.44) or patient-physician relationship (95% CI = 0.75	Pos	(self-constructed and recent and past interview questions) (p = .028) discussions of prognosis or life expectancy No changes in illness understanding in groups	reporting only past (p = .221) or no (p = .885) discussions of monunesis on life award-mon	Changes in perceived strength No associations or incorporation of patient-oncologist relationship coded prognostic	3 months arter patients onclodest reachonship visit, categorised in therepeutic (THC, pp. 109; PEPPL, p = .84)	Pos	audine (J = .029) room baseline to 3 months No associations	Detween coded progression prognostic discussion and changes in perceived	confidence in obtaining information and attention confidence in obtaining information and attention confidence in the confidence in the confidence of the co	or physicians (p = .5), nor between patient-reported prognostic discussion
Assessment of predictor		Cha	last or past visits (self-constructed interview int questions)		Patient-rated presence of discussion of life expectancy of	prognostic discussions 3 i					
Authors Assessm (year, country)		A.S. Epstein et al. Patient- (2016, USA) disc. life e	last (self ques		J.J. Fenton et al. Patient- (2018, USA) disc.	prog durir	infor				

	Relevant moderating effects ³	Ā	Age was a predictor of rating the expert/positive/collaborative and empathir approach as hopeful (more hope following expert/positive/collaborative and empathir approach among older vs. younger patients, p = .04 and p = .002) Anxiety was a predictor of rating the expert/positive/collaborative and empathic approach as hopeful (more hope following expert/positive/collaborative and empathic approach as hopeful, nor between age or anxiety and rating the avoidant approach as hopeful, nor between sex, religiosity, language, expected sunvival, time since metastatic diagnosis or involvement preferences and rating the expected sunvival, time since metastatic diagnosis or involvement preferences and rating the expected sunvival, time since metastatic diagnosis or involvement preferences and rating the expected sunvival, time since metastatic diagnosis or involvement preferences and rating the expected sunvival, time since metastatic diagnosis or involvement preferences	Education was a predictor of agreeing that the best case scenario conveyed hope (more agreement among higher vs. lower educated
	Relevant main effects ²	patient-oncologist relationship (THC, p = .21, P EPPT, p = .13) from baseline to 3 months baseline to 3 months Positive associations between discussions of life expectancy or prognosis and terminal illness acknowledgement (p = .036) and a trend for recognition of incurable disease (p = .085) controlled for gender differences No associations between discussions of life expectancy or prognosis and knowledge of advanced disease stage (p = .506) controlled for gender differences	Rated as most hopeful communication behaviours were offering the most up-to-date treatment (90%), appearing to know all there is to know about the patient's cancer (87%), occasional use of humour (80%), telling that the pain will be controlled (87%) and telling that the pain will be controlled (87%) and telling all treatment oppions (83%), shown in univariate analyses. Rated as not hopeful were appearing to be nervous or uncomfortable (91%), giving prognosis to family first (87%), use of euphenisms (82%), avoiding talking about cancer and only discussing treatment (75%) and giving good news first and then bad news (72%) Giving statistics about life expectancy was rated evenly hopeful (30%), not hopeful (30%) and neutral (30%) and neutral (30%) in univariate analyses, just like expressing uncertainty about the disease course (35% vs. 30% vs. 35%).	More patients agreed that explaining 3 scenarios vs. median survival would make sense (93% vs. 76%), be helpful (93% vs. 76%), help
	Patient outcome and assessment	Changes in illness understanding from pre- to post-scan visit, categorised in the acknowledgement of terminal disease, incurable disease and disease stage (self-constructed interview questions)	Hopefulness (survey questions based on Buttow et al. [52] and Sardell and Trierweiler [51])	Attitudes to different types of prognostic information categorised in making sense, being helpful, being helpful to make plans for the future.
(par	Assessment of predictor	Patient-rated presence of discussion of prognosis or life expectancy during the last or past visits (self-constructed interview questions)	Providing communication behaviours of physicians that might convey or discourage hope (written vignettes)	Providing the shottest 5–10%, middle 50% and longest 5–10%, of survival time or estimated median
Table 1. (Continued)	Authors (year, country)	K. Fletcher et al. (2013, USA)	R.G. Hagerty et al. (2005, Australia)	B.E. Kiely et al. (2013, Australia)

	Relevant moderating effects³	should be included (more agreement among higher vs. lower educated patients, p = .001) in bivariate analyses Education was a predictor of agreement among that the typical scenarios should be included (more agreement among that the typical scenarios should be included (more agreement among higher vs. lower educated patients, p = .05) and 3 scenarios were helpful (more agreement among higher vs. lower educated patients, p = .05) but not of agreeing that 3 scenarios made sense, were easy to understand (p = .50) in bivariate analyses (Cancer type was a a predictor of agreeing that 3 scenarios were reassuring (less agreement among breast cancer vs. other types, p = .009) and upsetting (more agreement among breast cancer vs. other types, p = .009) and upsetting (more agreement among females vs. males, p = .009) and upsetting (more agreement among females vs. males, p = .002) and worst case scenario was upsetting (more agreement among females vs. males, p = .002) in multivariate analyses (Time since exacter diagnosed > 1 year of agreement among patients of agreement among patients ≤ 60 vs. z 70 years, p = .001; more agreement among patients of agreement among patients since accorned news a upsetting (more agreement among patients since accorned news to p = .006; more agreement among patients since accorned news to p = .006; more agreement among patients of agreement among agreement among agreement among agreement among agreement among agreem	N A
	Relevant main effects ²	making plans for the future (88% vs. 70%), help family and cares (91% vs. 71%), convey hope (88% vs. 44%), be reassuring (60% vs. 40%), decrease anxiety (43% vs. 32%) and improve understanding of survival time (93% vs. 75%) (all $p < .001$) in bivariate analyses Fewer patients agreed that explaining 3 scenarios vs. median survival would take away hope (11% vs. 24%), be upsetting (24% vs. 36%) and increase arrivety (22% vs. 36%) and increase arrivety (22% vs. 35%) and increase arrivety (22% vs. 35%) (all $p < .001$) in bivariate	Less uncertainty ($p = .032$) and more satisfaction
	Patient outcome and assessment	helping family members and carers, conveying hope, taking hope away, being reassuring, being upsetting, decreasing anxiety, increasing anxiety, improving of understanding of survival time and preference to be included when explaining life expectancy (self-constructed survey questions)	Uncertainty and self-efficacy (survey questions based on Van Vliet
ntinued)	Assessment of predictor	(written vignettes)	Providing numbers or not (e.g. 2-year survival rate)
Table 1. (Continued)	Authors (year,		M. Mori et al. (2019, Japan)

anthropous turno	Relevant moderating effects ³		NA A
Dalmet many afficient	relevant main errects-	(p = .010) with more vs. less explicitness in bivariate analyses. No differences in anxiety (p = .198), self-efficacy (p = .279) or willingness to distouss advance care planning (p = .240) with more vs. less explicitness in bivariate analyses. No associations between explicitness and uncertainty (p = .074), satisfaction (p = .78), anxiety (p = .461), self-efficacy (p = .432) or willingness to discuss advance care planning (p = .371) anxiety (p = .461).	Less physical, behavioural and concentration-related symptoms of anxiety in patients (p = .024.0) with disclosure ws. (p = .024.0) with disclosure of incurability (p s0201) and family (p s0201) and family (g vC) in bivariate analyses Better recognition of disease condition in patients (p < .0001), as well as better communication among medical staff (between patient, family and between patient, family and medical staff (all p < .0001), with disclosure vs. non-disclosure of incurability (g vs. C) in bivariate analyses Better recognition of disease condition in patients (p < .0001) and family (p < .0001), as well as better communication among medical staff (all p < .0001), as well as better communication among medical staff (all p < .0001), with more vs. less specificity (C vs. D) in bivariate analyses of the symptoms, or pain or other symptoms, or pain physical symptoms, or pain physical symptoms, or pain specifically, in patients (p = .0001).
Dations on concentration	ratient outcome and assessment	et al. [54, 55]); satisfaction ((9SQ); anviety (STAI-state); willingness to discuss advance care planning (self-constructed survey question)	Health care provider perception of quality of terminal care categoriscal psychological state, recognition of disease condition, communication and physical symptoms (S1AS-J)
nued)	Assessment of predictor	during a simulated bad news consultation (video-recorded vignettes ⁵)	Researcher-rated level of prognostic communication categorised in (A) non-disclosure of carrier diagnosis; (B) disclosure of direct figanosis; (C) disclosure of fire-threatening diagnosis, e.g. metastasis and incurabitity, and (D) additional disclosure of poor prognosis, e.g. life expectancy (registered in medical records)
lable 1. (continued)	Authors (year, country)		N. Nakajima et al. (2012, Japan)

	rating																																							
	Relevant moderating effects³		V A											NA												NA														
	Relevant main effects ²	or family (p = .1892) between levels of prognostic commication in hivariahe analyses	No associations between statements of	optimism in fotal (p = .198) and about the past/present	pessimism about the	past/present $(p = .198)$ or uncertainty $(p = .988)$ and	patient-physician	concordance about chance of cure in bivariate analyses	Positive association between	statements of pessimism in total ($p = .006$) and	about the future $(p=.017)$	and patient-physician concordance about chance	of cure in multivariate	And More distress in patients with requested	disclosure vs. non-disclosure	of cure rates and/or ute expectancy $(v = .009)$, but	no differences in	confusion, anxiety, insecurity,	cominence, recung overwhermed or satisfaction (p. values	NA) in bivariate analyses	No differences in	emotional response or caticfaction	(p values NA)	between requested qualitative vs.	qualitative and quantitative information in bivariate analyses	Skin conductance level increased	(p = .004) in the affective and standard condition in bivariate	analyses, during disclosure of	incurability before the	manipulation started	communication about life	expectancy and treatment	options, there was better recall of information	(p = .035), more	perceived doctor	(p = .002) and reassurance	(p = .02), as well as a trend for	doctor emparing (p = .08), in the affective vs. standard	condition in bivariate analyses	Stronger decrease of skin conductance level $(p < .0001)$
	Patient outcome and assessment		Physician-patient concordance about chance of cure	based on physician and patient perception of cure	(self-constructed	survey question)								Satisfaction and emotional	response categorised in	retiet/distress, claritv/confusion.	reduced/increased anxiety,	security/insecurity,	streng triened/weakened confidence and feeling	supported/overwhelmed	(self-constructed survey	questions)				Skin conductance level	(microsiemens from the RTOPAC MP150): recall	(self-constructed survey	questions); non-abandonment,	reassurance of support and	QUOTE-COM)									
ned)	Assessment of predictor		Coding of optimistic, pessimistic or uncertain statements	about the past, present or future during	of test results, treatment or	prognosis (frequency counts)								Providing cure rates, cure rates	and life expectancy, or none,	and providing additional quantitative as well as	qualitative information (e.g.	5-year survival rate or	neulan surnivat unie) of not (individually adapted	medical consultation)						Providing reassurance or not	during a simulated bad news consultation	(video-recorded vignettes ⁵)												
Table 1. (Continued)	Authors (year, country)		T.M. Robinson et al. (2008, USA)											T. Rumpold et al.	(2015, Austria)											M.S.C. Sep et al.	(2014, The Netherlands)													

	Relevant moderating effects ³		A	Monitoring coping style was a moderator of the association between explicitness and uncertainty, anxiety, self-efficacy and satisfaction (more uncertainty and anxiety and less self-efficacy and astisfaction following explicitness among high monitors vs. low monitors, p = .007, p = .007, p = .008) No moderating effect of blunting coping style on the association between explicitness and patient outcomes, nor an effect of monitoring or blunting coping style on the association between effect of monitoring or blunting coping style on the association between effect of monitoring or blunting coping style on the association and patient outcomes (p values reported)	N N
	Relevant main effects ²	in the affective vs. standard condition, as well as standard condition, as well as an association between skin conductance level, and recall (p = .0.1) only in the affective condition, affect the manipulation in multiviorate analyses	Negative association between frequency of prognostic conversations and depressive symptoms (p < 01) No association between frequency of prognostic conversations	Negative associations between explicitness and uncertainty (p < .001). but not anxiety (p = .562). Positive associations between explicitness and self-efffacty (p = .004) and satisfaction (p < .001) Negative associations between affective communication and uncertainty (p = .001) Positive associations between affective communication and anxiety (p = .001) Positive associations between affective communication and self-efficacy (p < .001) and satisfaction (p ≤ .001) and satisfaction (p ≤ .001) and satisfaction for high explicitness and affective communication (all outcomes to communication (all outcomes of .001) and satisfaction for high explicitness and affective communication (all outcomes)	Better prognostic understanding in patients with vs. without prognostic discussion (\$P < 0.01) among the totals sample in bivariate analyses and among non-cancer patients and cancer patients separately (\$P\$ values NA) in subgroup analyses Nore discussion of care preferences with family (\$P < .01), as well as a trend for these discussions with providers (\$P < .10\$), in patients with vs. without prognostic discussion among the total sample in bivariate analyses No association between prognostic
	Patient outcome and assessment		Anxiety and depression (HADS)	Uncertainty and self-efficacy (self-constructed survey questions); anxiety (STAL-state); satisfaction (PSQ)	Prognostic understanding (interview question based on Quirt et al. [9]); discussion of care preferences with provider and family and documentation in a living will (self-constructed interview questions)
(pan	Assessment of predictor		Patient-rated frequency of prognostic conversations (PTPQ single item)	Providing numbers or not (e.g. 2-year survival rate) and providing reassurance or not during a simulated bad news consultation (video-recorded vignettes ³)	Patient-rated presence of prognostic discussion (interview questions based on Quint et al. [9])
Table 1. (Continued)	Authors (year, country)		J.A. Shin et al. (2016, USA)	L.M. Van Viiet et al. (2013, The Netherlands)	G.J. Wagner et al. (2010, USA)

	ıting		
	Relevant moderating effects ³		
	Relevant main effects ²	discussion and discussion of care preferences with family ($\rho = .21$) or providers ($\rho = .73$), or documentation of care preferences in a living will ($\rho = .18$) among the total sample in multivariate	analyses
	Patient outcome and assessment		
ed)	Assessment of predictor		
Table 1. (Continu	Authors As (year, country)		

CI confidence interval, ECG electrocardiogram, HADS Hospital Anxiety and Depression Scale, MQOL McGill Quality of Life Questionnaire, NA not available, PEPPI Perceived Efficacy in Patient-Physician Interactions scale, PSQ Patient Satisfaction Questionnaire, PTCC Prognostic and Treatment Choices scale, PTPQ Prognosis Treatment and Perceptions Questionnaire, QUOTE-COM Quality of Care Through the Patient's Eyes, RCT randomised controlled trial, SCID-5 Structured Clinical Interview for DSM-5, STAI-state State-Trait Anxiety Inventory state version, STAS-J Support Team Assessment Schedule–Japanese, THC The Human Connection scale, vs. versus

¹Prognosis as predictor was not further defined

Results of multivariate analyses were reported unless otherwise specified

Moderating effects encompassed interaction effects and predictors of patients' reactions to manipulated prognostic messages

Advance directives encompassed a living will, health care proxies and durable power of attorney for health care forms [56]

⁵Video-recorded vignettes used comparable scripts based on one previous qualitative study [55], all including disclosure of incurability before manipulation with explicit (life expectancy) and/or affective (life expectancy and/or treatment options) communication. Van Vliet et al. [54] and Mori et al. [57] examined the effect of the manipulation only. Sep Time domain parameters from heart rate variability series were the mean value of inter-beat intervals and the standard deviation. The median value along time was considered for further analysis. Frequency domain parameters from heart rate variability series were the median power spectral density of 5-s moving windows within two bandwidths: lowfrequency (from 0.04 to 0.15 Hz) and high-frequency (from 0.15 to 0.4 Hz) bandwidths. low-frequency reflects sympathetic activity with some degree of parasympathetic measure, et al. [53] and Danzi et al. [58] reported on of the effect of disclosing incurability additionally

and high-frequency derives from vagal or parasympathetic activity. The ratio between low-frequency and high-frequency is regarded as an index of sympathetic-parasympathetic

balance on heart rate modulation and reflects the sympathovagal interaction of the autonomic nervous system [58]

Quality assessment

Considering the twelve non-experimental studies, the definition of prognosis, sampling strategy, use of validated tools for patient outcomes, statistical tests and controlling for confounders were often satisfactory. However, a justified and satisfactory sample size, and comparability of responders and non-responders, or of responders with the target population, was frequently lacking. Hence, there might be problems with power and selection bias. Eight non-experimental papers attained $\geq 50\%$ of the maximum score. Four studies had relatively low quality (Table 2 in Appendix 3).

The manipulation of the predictor was well described in all six experimental studies; the definition of prognosis was clear in most. Additionally, randomisation, allocation, blinding of patients, comparison of groups and presence of equivalent conditions were adequate in both of the controlled trials. Blinding of the data analyst was not done in any of the experiments, and most lacked comparison of responders and non-responders, or comparison with the target population. Altogether, six papers showed methodological quality of \geq 50% (Table 3 in Appendix 4).

Prognostic disclosure

The effect of disclosing prognosis on advanced cancer patients was investigated by thirteen studies. Patient outcomes were categorised into information-related outcomes (e.g. understanding of prognosis), physical outcomes (e.g. symptoms), physiological outcomes (e.g. arousal), psychological outcomes (e.g. depression), relational outcomes (e.g. physician-patient relationship) or care preferences (e.g. documentation of treatment choices).

Information-related outcomes

Five studies assessed the association between disclosure of incurability and/or life expectancy and patients' prognostic understanding. All papers demonstrated significant positive associations [22, 59–62], although two examined a partially similar sample [59, 62]. Furthermore, a significant negative association between disclosure of life expectancy and length of patients' life expectancy self-estimates, and a positive association with the accuracy of these estimates, was found [22].

Physical outcomes

One study investigated the association between disclosure of incurability and physical symptoms, such as pain, and did not show a significant difference [60].

Physiological outcomes

Two studies examined the association between disclosure of incurability and physiological outcomes. In the experiment of Sep et al. [53], a significant increase of skin conductance was detected after physicians revealed the absence of cure in a simulated bad news consultation. In a similar experiment, Danzi et al. [58] studied activation of the autonomic nervous system through cardiac measures. Findings suggested significantly reduced parasympathetic activity, as well as increased sympathetic activity, among a part of the sample following

disclosure of incurability [58]. Increased physiological arousal, revealed in both studies, can be regarded as a marker of emotional stress [53, 58].

Psychological outcomes

Five studies investigated the association between prognostic disclosure and psychological outcomes, of which two had relatively low quality (< 50%) [63, 64]. Cripe et al. [63] discovered a significant negative association of patientrated extent of discussions about life expectancy with depressive symptoms. Moreover, depressive symptoms were less prevalent among respondents perceiving a worse prognosis when a full (versus no or a brief) discussion about life expectancy had taken place [63]. Similarly, more conversations about the likely disease outcome were significantly associated with less depressive symptoms [65]. Regarding anxiety, significantly lower levels were found in patients and their family when incurability was disclosed [60]. Other findings did not show differences in depression [22, 63], anxiety [22, 63–65] or feelings of confusion, insecurity, support or confidence in relation to prognostic discussions [64]. In contrast with previous findings, Cripe et al. [63] additionally revealed significantly more anxiety among respondents who perceived a worse prognosis after a full (versus no or a brief) discussion about life expectancy. Another study discovered more distress in patients who requested cure rates and survival estimates during an individually adapted consultation compared with patients who did not [64].

Relational outcomes

Four studies examined the association of prognostic conversations with relational outcomes. One showed a significant positive association between the disclosure of incurability and communication between patients, family and professionals [60]. Another study demonstrated significantly increased therapeutic alliance 3 months after discussions about incurability and survival [66]. Therapeutic alliance did not change from baseline to 2 days to 7 days after the consultation, nor did patients' confidence in obtaining information and attention of their physicians at any time point [66]. A third paper revealed no significant association between conversations about life expectancy and the physician-patient relationship [22]. Finally, satisfaction with an individually adapted consultation did not differ between patients receiving requested information about cure rates and life expectancy and patients who denied such information [64].

Care preferences

Three studies assessed the association between prognostic disclosure and care preferences. Bradley et al. [56] demonstrated that patients reporting conversations about life expectancy were more likely to have had physician-patient discussions about life-sustaining treatment preferences. In a sample mainly consisting of terminal cancer patients, a significant association between the disclosure of patients' life-limiting illness and discussing care preferences with family, as well as a positive trend for this discussion with physicians, was observed in bivariate analyses only [61]. One study additionally revealed a significant association between disclosure and the documentation of a do-not-resuscitate order [56]. Another paper, with relatively low quality, could not confirm this result, nor were

significant differences found in length of the last admission before death or the administration of sedation near death [67]. No significant associations between prognostic disclosure and documentation of a living will, health care proxies or durable power of attorney for health care were reported either [56, 61].

Disclosure strategies

Different strategies for prognostic communication, potentially influencing the effect of disclosure, were investigated by nine studies. Disclosure strategies were categorised into the provision of explicit prognostic information, framing of prognostic information, affective communication and general communication behaviours. Patient outcomes were categorised as previously reported.

Explicitness of prognostic information

Five papers assessed the association between explicit prognostic information and patient outcomes. Explicit communication encompassed disclosure of more specific prognostic information (e.g. life expectancy in addition to incurability) [60] or provision of quantitative (e.g. survival rate, median survival time and/or range) instead of, or supplemental to, qualitative survival information [54, 57, 64, 68].

Considering information-related outcomes, one cross-sectional study noted significantly better recognition of disease condition among patients who received more specific prognostic information [60]. Concerning physical outcomes, specific prognostic information was not associated with physical symptoms in general, nor with pain or any other specific symptoms [60]. Regarding psychological outcomes, uncertainty was significantly lower after the communication of quantitative versus qualitative information (in bivariate [57] and multivariate [54] analyses). Van Vliet and colleagues [54] additionally revealed enhanced self-efficacy regarding patients' ability to deal with the future following quantitative survival estimates, although Mori et al. [57] could not detect a similar effect. None of the included studies discovered a significant impact of explicitness on anxiety, distress, confusion, confidence, insecurity or feeling overwhelmed [54, 57, 60, 64]. Advanced cancer patients rated the provision of statistics about life expectancy evenly hopeful, neutral and not hopeful [68]. In terms of relational outcomes, communication between patients, family and professionals was significantly better for patients receiving more specific prognostic information [60]. Furthermore, two experimental studies demonstrated more satisfaction with communication of quantitative instead of qualitative survival estimates [54, 57]. Another paper reported no significant difference in patients' satisfaction with requested qualitative or supplemental quantitative information [64]. Lastly, studying care preferences, explicitness did not affect patients' willingness to discuss advance care planning with their oncologist [57].

Framing of prognostic information

Two experimental studies assessed the effect of framing prognostic information on patient outcomes by incorporating pessimistic, neutral or optimistic

statements [69] or using the worst, typical and best case scenario to explain life expectancy [70].

Concerning *information-related outcomes*, one study showed that patients were generally more optimistic about their prognosis than their physicians but were more likely to agree with their oncologist's estimated chance of cure when physicians made at least one statement of pessimism about the disease outcome [69]. Statements of optimism and uncertainty did not influence physician-patient concordance about the estimated chance of cure [69]. Furthermore, significantly more patients agreed that using the worst, typical and best case scenario to explain life expectancy, versus communicating median survival time, would improve their prognostic understanding, make sense, be helpful, help family and carers and help in making plans for the future [70]. In terms of *psychological outcomes*, significantly more patients agreed that presenting multiple survival scenarios would be reassuring, hopeful and less upsetting and would decrease anxiety, compared with median survival time [70].

Affective prognostic communication

Three experimental studies examined the impact of affective prognostic communication on patient outcomes [53, 54, 58]. The effect of affective communication was investigated by comparing respondents' reactions to video-recorded simulated bad news consultations, including communication about survival and treatment options, with and without physicians' reassurance of non-abandonment [54] and support [53, 58]. The absence of affective messages was referred to as standard communication [53, 54, 58].

Concerning information-related outcomes, one randomised controlled trial discovered better recall during affective versus standard communication [53], while another could not find a similar effect on self-perceived recall ability or actual recall [58]. Regarding physiological outcomes, affective communication led to a significantly stronger decrease of physiological arousal compared with standard communication, based on skin conductance levels [53]. Findings on cardiac measures varied, suggesting more sympathetic as well as parasympathetic activity of the autonomic nervous system during standard versus affective communication [58]. Combining information-related and physiological outcomes, both trials investigated whether the association of physiological arousal with recall differed between the affective and standard communication groups. Based on decreased skin conductance levels [53] and increased parasympathetic activity [58], reduced physiological arousal only led to improved recall during affective communication. However, based on increased heart rate variability, reduced physiological arousal only led to improved recall during standard instead of affective communication [58]. In terms of psychological outcomes, anxiety and uncertainty scores were significantly lower following physicians' reassurance of non-abandonment, while self-efficacy was significantly higher compared with standard communication [54]. Lastly, relational outcomes reflected participants' perceptions of the oncologist's behaviour in the simulated consultation. Affective communication, as compared to standard communication, led to significantly higher rates of satisfaction, perceived physician's support, non-abandonment, reassurance and empathy, but not engagement with the patient [53, 54, 58].

General communication behaviours

Hagerty and colleagues [68] investigated wide-ranging communication behaviours of physicians, which might influence hope during prognostic discussions. Physicians offering the most up-to-date treatment, appearing to know everything about a patient's cancer, using humour occasionally, telling pain will be controlled and communicating all treatment options were rated as most hope giving by advanced cancer patients. Those communication behaviours were labelled as expert, positive and collaborative approaches. In contrast, physicians appearing to be nervous, giving prognosis to family first, using euphemisms, avoiding talking about cancer, only discussing treatment and giving good news before bad news were not perceived as hope conveying in this experimental study. Those behaviours were labelled avoidant. Similar to providing survival statistics, physicians' expression of uncertainty about the disease course was rated evenly hopeful, neutral and not hopeful by metastatic cancer patients [68].

Moderating factors

Factors moderating the effect of disclosure strategies on patient outcomes were investigated by three studies. Considering explicit prognostic communication, patients with a strong monitoring coping style (i.e. often seeking detailed information) were significantly more anxious and uncertain and less self-efficacious and satisfied after explicitness than patients with lower monitoring scores [54]. A blunting coping style (i.e. often avoiding information) did not alter the effect of explicit survival information on patient outcomes [54]. With respect to framing prognostic information, higher educated patients were significantly more likely to agree that a typical case scenario, as well as a best case scenario, should be included when communicating life expectancy. Higher education was also associated with agreeing that the best case scenario conveyed hope, and that explaining three scenarios was helpful. No associations were found between education and agreeing that all scenarios were easy to understand [70]. According to the same study, females were significantly more likely to agree that all survival scenarios would help in making plans for the future, and that explaining a worst case scenario was upsetting. Being diagnosed with cancer more than 1 year ago and age younger than 70 were associated with the latter opinion as well. Furthermore, breast cancer patients were significantly less likely to agree that communicating a typical, best and worst case scenario was reassuring, as compared to patients with different primary tumour sites. Instead they were more likely to find this strategy upsetting [70]. Finally, concerning general communication behaviours, more anxious patients rated expert, positive, collaborative and empathic approaches to communicate prognosis (e.g. appearing to know everything, working as a team, expressing feelings) as significantly more hope conveying than less anxious patients [68]. The same result was found for older versus younger patients [68].

Discussion

Main findings

Prognostic disclosure

Integrating the results of all included studies examining prognostic disclosure, and considering methodological quality, evidence shows an association with improved prognostic awareness [22, 59-62]. This suggests that the previously mentioned high rate of prognostic unawareness [13-15] could reasonably be addressed by physicians through prognostic discussions. Moreover, Yun and colleagues [28] illustrate that it might be important for physicians to inform patients of their terminal status, as patients who became aware by their worsening condition or by chance reported lower quality of life.

Addressing oncologists' worries about harming patients through disclosure [6, 38, 39], the available evidence does not seem to confirm or deny this concern indisputably. Based on two experiments of satisfactory quality, disclosing incurability likely increases immediate physiological arousal [53, 58], indicating that the confrontation with prognostic information affects patients emotionally in that very moment. Literature examining the association of disclosure with longer-term psychological outcomes is limited and of varying quality. Furthermore, these findings are mixed, for which explanations are not found. Taken with caution, however, most point to either no or positive associations between prognostic disclosure and psychological well-being [22, 60, 63–65], which might be reassuring for physicians.

The reported discrepancy between physiological indices of emotional stress and psychological self-report measures could reflect the difference between immediate and delayed responses. Still, it should be taken into consideration that existing measures may not be sensitive enough to capture the complex emotional consequences of prognostic disclosure. Moreover, the effect of prognostic communication is often examined at group level, possibly disregarding subgroups with different reactions.

Accounting for physicians' worries about the physician-patient relationship [6, 38, 39], two papers show no or positive associations with communicating life expectancy [22, 66]. Additionally, three studies hint to a potential positive association between disclosure and physician-patient communication in general [60] or the discussion of care preferences specifically [56, 61]. Hence, these articles of satisfactory quality suggest that prognostic communication could strengthen the relation and stimulate patients' role in decision-making [61]. Sufficient evidence for an association between disclosure and the documentation of care preferences, actual care provided or physical outcomes could not be established [56, 60, 61, 67]. Remarkably, even though a key argument for prognostic disclosure is informed decision-making, very few papers investigated the association between prognostic communication and treatment decision-making.

Disclosure strategies

The limited number of studies, variety of examined strategies and hypothetical nature of the included experiments complicate drawing conclusions about the effect of different disclosure strategies. Hence, further research is needed to formulate strong recommendations. Based on the existing literature, we tentatively identify four approaches to prognostic communication.

The first strategy encompasses physicians' provision of more explicit, rather than imprecise, prognostic information. Three studies of satisfactory (and one with lower) quality suggest either no or a beneficial effect of explicitness on patients' psychological well-being and satisfaction with the consultation [54, 57, 60, 64]. One indicates improved recognition of disease condition [60]. According to the broader literature, being transparent about the difficulty of formulating individual survival estimates is considered helpful when communicating prognosis [44, 54]. Nonetheless, individual differences should be acknowledged, as another paper shows equally large proportions of respondents rating the provision of statistics and communication of uncertainty hopeful, neutral and not hopeful [68].

Second, physicians' tendency to stress the best case scenario, possibly inducing patients' overestimation of life expectancy [34, 38, 40, 41], could be complemented with a typical and worst case scenario. Both multiple survival scenarios and pessimistic statements improve patients' prognostic awareness, according to two studies of satisfactory quality [69, 70]. This realistic strategy might tackle oncologists' fear of leading patients to focus on a single number [34, 39], help patients to hope for the best, but prepare for the worst [34], and may prevent troublesome consequences of prognostic unawareness [16, 17, 22, 29–33].

A third disclosure strategy involves physicians' reassurance about non-abandonment and support while communicating life expectancy, investigated by three experiments of satisfactory quality. Findings hint to a beneficial effect of affective communication on patients' physiological arousal, psychological well-being, satisfaction with the consultation and most measures of perceived physician's support [53, 54, 58]. The effect of affective communication on recall of information, as well as the joint relation with physiological arousal, remains inconclusive [53, 58]. Nonetheless, this approach should satisfy patients' need to be looked after and allow for a sense of hope [54].

Fourth, hope might be fostered by physicians through expert, positive and collaborative behaviour during prognostic communication [68]. Incorporating wider research, oncologists are advised to address other sources of hope as well, rather than just medical information, like faith, inner peace, dignity, meaningful life events, relationships or humour [44, 54].

The combination of explicit and affective communication, multiple survival scenarios and expert, positive, collaborative behaviour might be most promising, although more scientific support is needed.

Moderating factors

Substantial evidence to identify patient and context characteristics that determine individual reactions to prognostic communication is limited. One study demonstrates that patients who tend to seek detailed information are more anxious and uncertain, and less self-efficacious and satisfied following

explicitness [54]. An explanation to this counterintuitive finding might be high monitors' generally lower satisfaction with information compared with patients showing a low monitoring coping style [54, 71, 72]. Another study suggests that higher educated patients are more likely to find explaining three survival scenarios helpful, and that breast cancer patients are more likely to find this strategy upsetting [70]. An obvious explanation for these findings was not provided, and the authors recommend communicating multiple scenarios to all advanced cancer patients preferring disclosure nevertheless [70]. Lastly, one paper indicates that more anxious and older patients particularly feel hopeful after prognostic communication displaying expertise, positivity, collaboration and empathy [68]. The latter is in line with earlier research, confirming that sensitive communication with emotional support is especially important for older cancer patients [73].

Limitations

This paper presents a novel attempt to offer comprehensive, evidence-based guidance for physicians' clinical practice by examining the independent effect of prognostic communication on advanced cancer patients. Despite the suggested disclosure strategies, some patients might prefer and fare better with prognostic ignorance [11, 74, 75]. Reasonably, some experts in the field make a case for prognostic silence, emphasising its protective function [76•, 77]. Exploring a patient's personal information preferences beforehand, indeed, is essential in prognostic conversations [44, 54]. Caregivers' information needs should be addressed additionally, as potential dissimilarities could influence prognostic discussions [44].

The reviewed studies have some limitations, which should be taken into account when interpreting the results. Firstly, prognostic disclosure is often assessed with self-report or medical record registration instead of direct observations, which might not reflect the actual extent of prognostic discussions [22, 59]. Secondly, the used study designs entail restrictions. Non-experimental study designs cannot imply causation [63, 66], whereas the controlled context of experiments diminishes the complexity of clinical interactions [54, 57]. Furthermore, respondents acting as analogue patients in experimental studies may react differently than actual patients [53, 54]. Still, since manipulation of prognostic communication in real-life settings is ethically unfeasible, we have to rely on the combination of these study types.

The current paper has shortcomings too. The authors solely searched literature indexed by PubMed and PsycInfo, possibly excluding relevant articles in other databases. Furthermore, attempts to facilitate prognostic communication by means of interventions are not discussed. Studies on advance care planning, early palliative care, decision aids, question prompt lists or communication skills training may provide supplemental information on the effect of prognostic messages.

Future directions

Considering the limits of literature to-date, future research should expand knowledge about the influence of different prognostic communication formats on patients to convey more pronounced advice, with a focus on psychological well-being over time and treatment decision-making. Studies should attempt to

overcome the one-size-fits-all approach, by exploring patients' individual information preferences and differences in reactions to prognostic communication. Knowledge of moderators, as well as enhanced prognostic prediction models, could assist oncologists in tailoring their messages. Finally, future research should optimise the development and implementation of communication interventions to put guidelines into practice and ultimately improve prognostic communication.

Conclusions

Altogether, this systematic review synthesised today's literature on the effect of prognostic communication on patient outcomes in palliative cancer care and moderators of this relation. Addressing the research aims, we conclude cautiously that, if preferred by patients, oncologists can disclose prognosis using sensible strategies. Displaying expertise, positivity and collaboration, while offering explicit prognostic information with multiple survival scenarios and reassurance of support, likely offers an inclusive approach physicians can rely on.

Opinion statement

Making more deliberate communicative choices starts with physicians' awareness of their propensity to deliver ambiguous messages. Based on current knowledge and ethical principles, we recommend oncologists to communicate prognosis to patients who wish to know, using a balanced approach. Allowing for preparation as well as hope, we advise mentioning the worst, typical and best case scenario of survival, instead of one number or the optimistic scenario only. Worries about harming patients by truth-telling should not prevent physicians from at least offering prognostic information, as these worries might also originate from personal fears or discomfort. We stimulate physicians to acknowledge patients' emotions, which often reflect a normal reaction to lifechanging messages, and may not necessarily persist. Additionally, we encourage physicians to support patients by reassuring non-abandonment and fostering different types of hope. Oncologists can incorporate the uncertainty of individual estimates by stressing the possible deviation from group-based survival information. Finally, we emphasise the need to individualise care. It is essential to be aware of patients' frame of reference and to recognise potential differences with one's own social, cultural and religious context. Discussing prognosis is considered an ongoing process. Therefore, individual information needs have to be explored timely, thoroughly and repeatedly. Subsequently tailoring to patients' preferences would be most advantageous.

Funding Information

This project was funded by the Dutch Cancer Society (KWF Kankerbestrijding, grant number 11547).

Compliance with Ethical Standards

Conflict of Interest

Naomi C. A. van der Velden declares that she has no conflict of interest.

Maartje C. Meijers declares that she has no conflict of interest.

Paul K. J. Han declares that he has no conflict of interest.

Hanneke W. M. van Laarhoven declares that she has no conflict of interest.

Ellen M. A. Smets declares that she has no conflict of interest.

Inge Henselmans declares that she has no conflict of interest.

Human and Animal Rights and Informed Consent

This article does not contain any studies with human or animal subjects performed by any of the authors.

Appendix 1. Search strategies

Search strategy PubMed

(Prognosis[Mesh] OR prognos*[tiab] OR prognos*[other term] OR "life expectancy" [Mesh] OR "life expectancy" [tiab] OR mortality [Mesh] OR "disease course"[tiab] OR "mortality risk"[tiab] OR death[Mesh] OR survival[Mesh] OR survival[tiab] OR "bad news"[tiab] OR "truth disclosure"[Mesh] OR "truth disclosure"[tiab]) AND (Communication[Mesh] OR communicat*[tiab] OR communicat*[other term] OR conversat*[tiab] OR messag*[tiab] OR "knowledge transfer"[tiab] OR informing[tiab] OR information[ti] OR "patient education as topic" [Mesh] OR "patient education" [tiab] OR "nonverbal communication" [Mesh] OR "health communication" [Mesh] OR "psychotherapeutic processes" [Mesh] OR discussi* [tiab] OR dialog* [tiab] OR "truth disclosure"[tiab] OR consult*[tiab] OR "professional-patient relations"[Mesh] OR "physician patient relations" [Mesh] OR (((((patient[tiab] OR client[tiab]))) AND ((physician[tiab] OR specialist[tiab] OR oncologist[tiab] OR professional[tiab] OR doctor[tiab] OR clinician[tiab] OR provider[tiab]))) AND ((interaction*[tiab] OR relation*[tiab] OR communication*[tiab] OR discussion*[tiab])))) OR (((((patient[Other Term] OR client[Other Term]))) AND ((physician[Other Term] OR specialist[Other Term] OR oncologist[Other Term OR professional Other Term OR doctor Other Term OR clinician Other Term] OR provider[Other Term]))) AND ((interaction*[Other Term] OR relation*[Other Term] OR communication*[Other Term] OR discussion*[Other Term])))) AND ((Neoplasms[Mesh] OR neoplasm*[tiab] OR "Medical Oncology" [Mesh] OR oncolog* [tiab] OR oncolog* [Other Term] OR cancer*[tiab] OR carcino*[tiab] OR tumor*[tiab] OR tumour*[tiab] OR sarcoma*[tiab] OR malignan*[tiab] AND "terminal care"[Mesh] OR terminal*[tiab] OR terminal*[Other Term] OR "palliative care"[Mesh] OR palliati*[tiab] OR palliati*[Other Term] OR Neoplasm metastasis[Mesh] OR metasta*[tiab] OR "disease progression"[tiab] OR "terminally ill"[Mesh] OR "advanced cancer"[tiab] OR "advanced cancer"[Other Term] OR "end of life"[tiab] OR "end-of-life"[tiab] OR "supportive care"[tiab] OR incurable[tiab] OR non-curable[tiab])) AND ((Humans[Mesh] AND English[lang] AND (aged, 80 and over[MeSH] OR aged[MeSH] OR middle age[MeSH] OR (middle

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age[MeSH] OR aged[MeSH]) OR adult[MeSH:noexp] OR adult[MeSH] OR young adult[MeSH])))) AND (Humans[Mesh] AND English[lang] AND (aged, 80 and over[MeSH] OR aged[MeSH] OR middle age[MeSH] OR (middle age[MeSH] OR aged[MeSH]) OR adult[MeSH:noexp] OR adult[MeSH] OR young adult[MeSH]))) Filters: Humans; English; 80 and over: 80+ years; Aged: 65+ years; Middle Aged: 45-64 years; Middle Aged + Aged: 45+ years; Adult: 19-44 years; Adult: 19+ years; Young Adult: 19-24 years

Search strategy PsycInfo

(((prognosis/ OR prognos*.ab,ti,id. OR "life expectancy"/ OR "life expectancy".ab,ti. OR "mortality risk"/ OR "mortality risk".ab,ti. OR "disease course"/ OR prediction/ OR severity/ OR "death and dying"/ OR survival.ab,ti. OR "bad news".ab,ti. OR "truth disclosure".ab,ti.) AND (communication/ OR communicat*.ab,ti,id. OR conversation/ OR conversat*.ab,ti. OR messages/ OR messag*.ab,ti. OR "knowledge transfer"/ OR "knowledge transfer".ab,ti. OR information/ OR informing.ab,ti. OR information.ti. OR "client education"/ OR "patient education".ab,ti. OR "interpersonal communication"/ OR "verbal communication"/ OR "nonverbal communication"/ OR "oral communication"/ OR "communication skills"/ OR "therapeutic processes"/ OR discussi*.ab,ti. OR dialog*.ab,ti. OR "truth disclosure".ab,ti. OR consult*.ab,ti. OR ((patient OR client) AND (physician OR specialist OR oncologist OR professional OR doctor OR clinician OR provider) AND (communication* OR discussion* OR interaction* OR relation*)).id,ti,ab.) AND (((exp neoplasms/ NOT "benign neoplasms"/) OR neoplasm*.ab,ti. OR oncology/ OR oncolog*.ab,ti,id. OR cancer*.ab,ti. OR carcino*.ab,ti. OR tumor*.ab,ti. OR tumour*.ab,ti. OR sarcoma*.ab,ti. OR malignan*.ab,ti.) AND ("terminal cancer"/ OR terminal*.ab,ti,id. OR "palliative care"/ OR palliati*.ab,ti,id. OR metastasis/ OR metasta*.ab,ti. OR "disease progression"/ OR "disease progression".ab,ti. OR "terminally ill patients"/ OR "advanced cancer".ab,ti,id. OR "end of life".ab,ti. OR "end-of-life".ab,ti. OR "supportive care".ab,ti. OR incurable.ab,ti. OR non-curable.ab,ti.))) AND (adulthood 18 yrs older OR aged 65 yrs older OR middle age 40 64 yrs OR thirties 30 39 yrs OR very old 85 yrs older OR young adulthood 18 29 yrs).ag. AND (dutch OR english).lg.)

Appendix 2. Quality assessment tools

Checklist for quality assessment of non-experimental studies

This scale has been adjusted from the adapted Newcastle-Ottawa Scale¹, to perform a study-specific quality assessment of non-experimental studies for the systematic review of Van der Velden et al. (The Effect of Prognostic Communication on Patient Outcomes in Palliative Cancer Care: a Systematic Review, 2020). A maximum score of 16 points can be attained.

Studies assessed using this scale are as follows: Aoki et al. (1997), Bradley et al. (2001), Cripe et al. (2012), Enzinger et al. (2015), Epstein et al. (2016), Fenton et al. (2018), Fletcher et al. (2013), Nakajima et al. (2012), Robinson et al. (2008), Rumpold et al. (2015), Shin et al. (2016) and Wagner et al. (2010).

- 1. Representativeness of the sample (0-2 points)¹
 - i. Truly representative of the average in the target population (i.e. all subjects or random sampling)**
 - ii. Somewhat representative of the average in the target population (i.e. non-random sampling)*
 - iii. Selected group of users (i.e. snowballing or convenience sampling)
 - iv. No description of the sampling strategy
 - 2. Sample size (0-2 points)¹
 - i. Justified (power analysis) and satisfactory**
 - ii. Not justified but likely satisfactory ($N \ge 128$, sample size required to test for a medium-sized mean difference between two independent groups, e.g. disclosure or non-disclosure)*
 - iii. Justified but not satisfactory, or not justified and likely not satisfactory (N < 128)
- 3. Non-responders (0–2 points)¹
 - i. Response rate is satisfactory (i.e. > 50%) and comparability² of responders' and non-responders' characteristics is established**
 - ii. Response rate is not satisfactory (i.e. < 50%), but comparability of responders' and non-responders' characteristics is established*
 - iii. Response rate is satisfactory (i.e. > 50%), but comparability of responders' and non-responders' characteristics is not established
 - iv. Response rate is not satisfactory (i.e. < 50%) and comparability of responders' and non-responders' characteristics is not established, or response rate is not reported

Comparability

- 4. Confounding factors (0–2 points)¹
 - i. The study controls for potential confounder(s)**
 - ii. No control for confounders

Exposure

- 5. Definition of the predictor (prognostic communication) (0–2 points)³
 - i. Definition of the predictor is clearly described **
 - ii. Unclear or no description of the definition of the predictor

- 6. Assessment of the predictor (prognostic communication) (0-2 points)³
 - i. Observation of real consultations (coding of audio-recorded visits)**
 - ii. Tailoring of real consultations or medical record registration*
 - iii. Self-report by doctors or patients of real consultations
 - iv. No description of assessment of the predictor

Outcome

- 7. Validation of the measurement tools for patient outcomes $(0-2 \text{ points})^1$
 - i. Validated measurement tools (self-report or structured interview) or medical record registration of DNRs**
 - ii. Non-validated measurement tools included (self-report, interview question or medical record registration of conversations), but the tools are all available or described*
 - iii. No description of measurement tools
- 8. Statistical test (0–2 points)¹
 - i. The statistical test used to analyse the data is clearly described and is appropriate (including multilevel analyses if possible and deemed necessary, and corrections for multiple testing when > 20 relations were tested), and all significant and non-significant results are reported including the measurement of association and p values or confidence intervals * *
 - ii. Two out of three of the above present*
 - iii. The statistical test is not appropriate, incomplete or not described

Total score

Checklist for quality assessment of experimental studies

This is a self-constructed checklist to perform a study-specific quality assessment of experimental studies for the systematic review of Van der Velden et al. (The Effect of Prognostic Communication on Patient Outcomes in Palliative Cancer Care: a Systematic Review, 2020), consisting of two parts. Experimental studies using a within-subjects design are assessed with part A. Experimental studies using a between-subjects design are assessed with parts A and B. Items of the

¹Adjusted from the adapted Newcastle-Ottawa Scale

²Comparability is considered established when responders have been compared with non-responders or with the total population, and no significant differences were found

³Self-constructed item specified to suit the research questions of the systematic review of Van der Velden et al. (The Effect of Prognostic Communication on Patient Outcomes in Palliative Cancer Care: a Systematic Review, 2020)

Cochrane Consumer and Communication Review Group criteria, the Cochrane Collaboration's tool for assessing risk of bias and the adapted Newcastle-Ottawa Scale were adjusted and used for the current checklist. A maximum score of 8 points can be attained using part A only. A maximum score of 13 points can be attained using parts A and B.

Studies assessed using this scale are as follows: Danzi et al. (2018), Hagerty et al. (2005), Kiely et al. (2013), Mori et al. (2019), Sep et al. (2014) and Van Vliet et al. (2013).

Part A

- 1. Was the definition of the predictor (prognostic communication) clearly described?1
- 2. Was the manipulation of the predictor (prognostic communication) clearly described?1
- 3. Was the data analyst blinded (i.e. was the code only broken after conclusions were drawn)?^{3, 6}
- 4. Were no outcome data missing, or were missing outcome data unlikely to be related to true outcome, balanced across conditions with similar reasons, assumed not to have a clinically relevant impact on the intervention effect or imputed with appropriate methods?³
- 5. Were validated tools (i.e. well-known questionnaires, reliable coding schemes or standardised instruments for physiological measurements) used to assess all outcomes that were related to prognostic communication or were non-validated tools at least all available or described?²
- 6. Was the sample size justified and satisfactory, or not justified but presumably satisfactory ($N \ge 128$, sample size required to test for a medium-sized mean difference between two independent groups, e.g. disclosure or non-disclosure)?⁵
- 7. Was the response rate satisfactory (i.e. > 50%) and was comparability of responders' and non-responders' characteristics established?^{5, 7}
- 8. Was the statistical test used to analyse the data clearly described and appropriate (including multilevel analyses if possible and deemed necessary, and corrections for multiple testing when > 20 relations were tested), and were all significant and non-significant results reported including the measurement of association and p values or confidence intervals?5

Part B

- 9. Was the method of randomisation adequate (i.e. truly random)?^{2, 6}
- 10. Was allocation concealed (i.e. could allocation to conditions have been influenced)?^{2, 6}

- 11. Was the (analogue) patient blinded (i.e. did the patient know to what condition s(he) was allocated)?²
- 12. Were groups compared at baseline on at least one potential determinant of the outcomes (i.e. socio-demographics, medical characteristics and primary outcomes) and were the analyses properly controlled for differences?²
- 13. With the exception of the trial intervention, were the experimental and control condition equivalent (i.e. was a placebo added to the control condition)?⁴

Total score

¹Self-constructed item specified to suit the research questions of the systematic review of Van der Velden et al. (The Effect of Prognostic Communication on Patient Outcomes in Palliative Cancer Care: a Systematic Review, 2020)

²Among the criteria formulated by the Cochrane Consumer and Communication Review Group

³Adjusted from the Cochrane Collaboration's tool for assessing risk of bias

⁴Adopted from Henselmans et al. [50]

⁵Adjusted from the adapted Newcastle-Ottawa Scale

⁶Considered fulfilled when called random, concealed or blinded by authors or when the text explicitly refers to such methods (such as mention of 'sealed opaque envelopes', a cover story for patients in the control group or separate consent forms)

⁷Comparability is considered established when responders have been compared with non-responders or with the total population, and no significant differences were found

Appendix 3

Table 2. Quality assessment of non-experimental studies

Authors (year)	Selection			Comparability	Exposure		Outcome		Total and % of
	(1) Representativeness of the sample	(2) Power and sample size	(3) Non- response	(4) Confounders	(5) Definition of prognosis	(6) Assessment of predictor	(7) Validated tools	(8) Statistics	maximum score
Aoki et al. (1997)	1	1	1	1	1	*	* *	*	4/16 = 25.0%
Bradley et al. (2001)	*	*	*	* *	*	* *	*	* *	13/16 = 81.3%
Cripe et al. (2012)	ı	1	ı	* *	* *	ı	* *	*	7/16 = 43.8%
Enzinger et al. (2015)	*	*	* *	* *	* *	ı	*	*	10/16 = 62.5%
Epstein et al. (2016)	*	* *	1	* *	* *	ı	*	*	9/16 = 56.3%
Fenton et al. (2018)	*	*	1	* *	*	* *	* *	* *	12/16 = 75.0%
Retcher et al. (2013)	*	1	1	* *	*	1	*	*	7/16 = 43.8%
Nakajima et al. (2012)	*	1	* *	* *	* *	*	* *	*	12/16 = 75.0%
Robinson et al. (2008)	*	*	1	* *	* *	* *	*	*	11/16 = 68.8%
Rumpold et al. (2015)	*	1	1	1	*	*	*	*	6/16 = 37.5%
Shin et al. (2016)	*	*	1	* *	*	ı	* *	* *	10/16 = 62.5%
Wagner et al. (2010)	*	*	ı	* *	* *	1	*	*	8/16 = 50.0%
A maximum of 2 points can be attained per item	ttained per item								

Appendix 4

Authors (year)	Part A							
Additions (year)	C 181							
	(1)	(2)	(3)	(4)	(5)	(9)	(7)	(8)
	Definition of	Manipulation	Blinded	Missing	Validated	Power and	Non-response	Statistics
	prognosis	of predictor	data	outcome	tools	sample		
			analyst	data		size		
Non-controlled experimental studies (within-subjects design)	udies (within-subjects design)							
Hagerty et al. (2005)	ı	*	ı	*	*	ı	*	1
Kiely et al. (2013)	*	*	1	*	*	*	ı	1
Mori et al. (2019)	*	*	1	*	*	*	ı	*
Van Vliet et al. (2013)	*	*	1	*	*	*	ı	*
Controlled experimental studies (between-subjects design)	(between-subjects design)							
Danzi et al. (2018)	ı	*	ı	*	ı	ı	ı	*
Sep et al. (2014)	*	*	ı	*	ı	1	1	*
	6						1	-
Autnors (year)	Part B						101	lotal and %
	(6)	(10)		(11)	(12)	(13)	of	of maximum
	Randomisation	Allocation		Blinded	Group	Equivalent		score
			Ď	patient	comparison	conditions	sus	
Non-controlled experimental studies (within-subjects design)	udies (within-subjects design)							
Hagerty et al. (2005)	NA	NA	NA	4	NA	NA	4	4/8 = 50.0%
Kiely et al. (2013)	NA	NA	NA	đ	NA	NA	5/	5/8 = 62.5%
Mori et al. (2019)	NA	NA	NA	4	NA	NA	/9	6/8 = 75.0%
Van Vliet et al. (2013)	NA	NA	NA	4	NA	N	/9	6/8 = 75.0%
Controlled experimental studies (between-subjects design)	(between-subjects design)							
Danzi et al. (2018)	*	*	*		*	*	8/1	8/13 = 61.5%
Sep et al. (2014)	*	*	*		*	*	9/1	9/13 = 69.2%
A maximum of 1 point can be attained per item	attained per item							

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