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Review Article

Interventions to Promote End-of-Life Conversations: A Systematic Review and Meta-Analysis

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Abstract

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Context. Although several interventions aimed to promote end-of-life conversations are available, it is unclear whether and how these affect delivery of end-of-life conversations. Measuring the processes associated with high-quality end-of-life care may trigger improvement.

Objectives. To estimate the effect of interventions aimed to promote end-of-life conversations in clinical encounters with patients with advanced chronic or terminal illness or their family, on process indicators of end-of-life conversations.

Methods. Systematic review with meta-analysis (PROSPERO no. CRD42021289471). Four databases (PubMed, CINAHL, PsycINFO, and Scopus) were searched up to September 30, 2021. The primary outcomes were any process indicators of end-of-life conversations. Results of pairwise meta-analyses were presented as Risk Ratio (RR) for occurrence, standardized mean difference (SMD) for quality and ratio of means (ROM) for duration. Meta-analysis was not performed when fewer than four studies were available.

Results. A total of 4,663 articles were scanned. Eighteen studies were included in the systematic review and 16 entered at least one meta-analysis: documented occurrence (n = 8), patient-reported occurrence (n = 4), patient-reported-quality (n = 4), duration (n = 4). There was significant variability in settings, patients' clinical conditions, and professionals. No significant effect of interventions on documented occurrence (RR 1.54, 95% CI 0.84–2.84; I² 91%), patient-reported occurrence (RR 1.52, 95% CI 0.80–2.91; I² 95%), patient-reported quality (SMD 0.83, 95% CI –1.06 to 2.71; I² 99%), or duration (ROM 1.20, 95% CI 0.95–1.51; I² 65%) of end-of-life conversations was found. Data on frequency were conflicting. Interventions targeting multiple stakeholders promoted earlier and more comprehensive conversations.

Conclusion. Heterogeneity was considerable, but findings suggest no significant effect of interventions on occurrence, patient-reported quality and duration of end-of-life conversations. Nevertheless, we found indications for interventions targeting multiple stakeholders to promote earlier and more comprehensive conversations. J Pain Symptom Manage 2023;000:e1—e34. © 2023 American Academy of Hospice and Palliative Medicine. Published by Elsevier Inc. All rights reserved.

Key Words

Clinical encounter, conversations, end of life, meta-analysis, systematic review, terminally ill

Key Message

This systematic review with meta-analysis found no significant effect on process indicators of interventions aimed to promote end-of-life conversations in clinical encounters with patients with advanced chronic or terminal illnesses or their family. Heterogeneity was considerable and further investigation into their implementation as a collaborative, person-centered effort is warranted.

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Introduction

High-quality end-of-life care relies on optimal treatment of symptoms and psychosocial, spiritual and existential support for both patient and family, ^{1,2} and on knowledge of patients' care preferences to deliver care aligned with their preferences. This requires effective and regular communication such as in advance care planning (ACP) conversations and shared decision-making. ^{1,2}

Engagement in end-of-life conversations by patients or family, measured in different ways,³ varies significantly: from 18%⁴ to 73%⁵ in cancer to 9.5% up to 70% in nursing home settings.^{6,7} Indeed, how end-of-life conversations are being defined varies across studies,⁸ and adopting a broad definition may be reasonable.

Over the last 30 years, investments have been made to develop resources, strategies and conceptual frameworks for improving end-of-life conversations. 9-13 However, social and healthcare professionals (hereafter professionals)-led end-of-life conversations are still suboptimal from the perspective of both patients and their family.¹⁴ Their measurement based on quality standards represents a milestone and indicators have been identified at the level of physical environment, process, and outcome. 15,16 In serious illnesses, the conversation process mediates goal-concordant care, 17 which constitutes a priority outcome in palliative care. 18 Therefore, measuring this process is critical to ensure high-quality conversations and improve the care provided. Process indicators offer objective and straightforward measures on how care is delivered and can be tracked and easily targeted in interventions aimed to improve the quality of care. 19,20 National consensus bodies recommend ameliorations in the way end-of-life conversations are delivered with respect to their occurrence, quality, timing, frequency, and duration. 21-24

Several international palliative care guidelines recommend accurate documentation of end-of-life conversations in the clinical record not limited to their occurrence.²⁵ Terminally-ill patients who report any end-of-life communication have better quality of death compared to patients who do not²⁶ and the quality of dying improves as the quality of conversations increase.²⁷ High-quality end-of-life communication has been associated with improved outcomes for patient and family, 4,8,28 while absent, late, or poor professionals-patient/family communication contributes to aggressive, lower-quality and patients' non-preferred treatment.^{3,29} The "Conversation ready" framework for improving end-of-life care assumes that the quality of documentation depends on the effectiveness and reliability of patients and families engagement, and therefore it can be employed as a basic process measure.³⁰

There is widespread support for the need to start end-of-life conversations early in the disease trajectory to give patients and family support, promote preparedness for the upcoming worsening, and offer the opportunity to raise questions with professionals to sustain understanding and involvement of patients in making care decisions. ²⁵ Unfortunately, conversations usually take place late in the disease trajectory with professionals newly involved in the patient's care, and difficult decisions are made in the heat of the moment. ⁵ One-third of conversations usually take place within one month of death. ³¹

A series of conversations may be needed to promote understanding and shared decisions by making information more digestible and allowing processing time.³² When end-of-life conversations are held on an ongoing basis, they strengthen trust and rapport, and favor the elicitation of personal values and goals and their revisit as the patient's conditions change.³³ Also, family satisfaction increases with frequency^{34,35} of conversations and more frequent contact is associated with not providing aggressive care.³⁶

Finally, it is recommended to arrange adequate time for conversation to allow patients and families to ask questions and express their views without feeling rushed into a decision. Family satisfaction increases when professionals take the time to listen, but end-of-life conversations are often brief.

Several interventions aimed to promote end-of-life conversations have been developed but there is no review summarizing evidence on their effects on how end-of-life conversations are delivered. Previous systematic reviews mainly examined effects on communication skills, 40,41 and on patients' or family caregivers' outcomes. 42 Measuring process indicators that may predict high-quality end-of-life care can support quality improvement efforts by identifying areas of amelioration. Therefore, this paper looks at the end-of-life conversations in terms of occurrence, quality, timing, frequency, and duration. 21–24

The interventions aimed to promote end-of-life conversations have been assessed in trials that varied from explanatory, examining effects in ideal circumstances, to pragmatic in usual care conditions. The type of trial is often not stated explicitly. Describing the trial approach informs the applicability of interventions aimed to promote end-of-life conversations.

Therefore, we systematically reviewed the literature and conducted meta-analyses with the primary objective to estimate the effect of interventions aimed to promote end-of-life conversations in clinical encounters with patients with advanced chronic or terminal illnesses or their family on occurrence, quality, timing, frequency, and duration of end-of-life conversations. The secondary objective was to describe trial approaches (explanatory vs. pragmatic).

Method

Design

A systematic review according to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines⁴⁴ (Table A1) was performed. The protocol was registered on 6 December 2021 (PROSPERO CRD42021289471).

Search Strategy and Selection Criteria

Searching PubMed and one giant database (e.g., Scopus, Embase) at minimum is recommended for systematic reviews. We searched four databases (PubMed, CINAHL, PsycINFO, and Scopus) on 30 September 2021 from inception. Searches employed controlled vocabularies and free terms, without temporal or language limits. Search strategies were adapted for each database (Appendix 2). The reference lists of included articles were screened manually to identify potentially relevant publications. Also, the PROSPERO register of systematic reviews was searched for ongoing and recently completed reviews.

In the present study, end of life conversation was defined as a clinical interaction engaging professionals and patients with advanced chronic or terminal illness, or their family, and aimed to promote patient and/or family understanding of illness progression or care options, or elicit patient's care preferences at the end of life. 13 Studies were included if they 1) covered at least one element of this definition; 2) involved adults with advanced, progressive, incurable conditions approaching the end of life 15 and/or their family. Studies involving only family caregivers were also eligible; 3) were intervention studies (i.e., pre post studies, clinical trials, controlled clinical trials, and randomized controlled trials) aimed to promote end-of-life conversations in clinical encounters, regardless the clinical setting and people targeted (i.e., patients, family caregivers, and/or professionals); 4) reported on the effect of interventions on at least one process indicator of end-of-life conversations (i.e., occurrence, quality, timing, frequency, or duration); and 5) were published in peer reviewed journals.

We excluded studies on interventions which targeted the general public or older people more generally (e.g., health campaign, public health initiatives, and policy), or were part of multi-faceted programs and the effect of the individual component of the intervention that related to end-of-life communication was not clearly recognizable and assessable.

As we focused on process outcomes, studies only assessing the impact of interventions on family-related (e.g., psychological distress) or patient-related outcomes (e.g., hospitalizations) were also excluded. Patient/family care outcomes represented additional

data that were extracted for descriptive purposes, when available.

Article Screening and Study Selection

S.G. and Y.A. independently screened titles and abstracts, removed duplicates, and reviewed the full text of potentially relevant articles, and discussed any disagreement or uncertainty regarding eligibility until reaching consensus.

Quality Assessment

S.G. and Y.A. assessed study quality using the Qual-Syst tool for quantitative research. ⁴⁶ This 14-item tool scored whether specific criteria were met ("fully" = 2, "partially" = 1, "not at all" = 0). Items not relevant for a particular study design were excluded from the summary score that was calculated for each paper and allowed for direct comparison of studies with different designs. Studies were classified as high (>0.9), moderate (0.7-0.9) or low quality (<0.7).

Applicability Assessment

Trial approaches vary from explanatory, examining effects in ideal circumstances, to pragmatic when applicability of the intervention in usual care across a range of settings is relevant. The approach becomes more explanatory as it moves away from usual practice towards ideal circumstances, offering insights in whether the intervention could work at all.

S.G. and Y.A. independently assessed applicability using the PRagmatic Explanatory Continuum Indicator Summary-2 (PRECIS-2) tool, ⁴³ which focuses on trial design choices and comprises nine domains: 1) eligibility criteria; 2) recruitment; 3) setting; 4) organization (expertise and resources to deliver the intervention); 5) flexibility-delivery of intervention; 6) flexibility-measures to ensure adherence to the intervention; 7) follow up; 8) primary outcome; and 9) primary analysis. Each domain was scored on a 5-point scale (1 = very explanatory to 5 = very pragmatic). With insufficient information, we assigned a score of three.

A mean score across studies for each domain and a mean applicability rating for each study were calculated. Domains and studies were classified as mostly explanatory (<4) or mostly pragmatic (≥4) according to the mean domain score and the mean applicability rating, respectively.

Data Extraction

Data were entered into a standardized spreadsheet under the following headings: study characteristics; sample characteristics; narrative summary of findings; and quantitative results (number and proportion of individuals with self-reported or documented occurrence of endof-life conversations in medical record; mean values (standard deviation [SD]) of patient-reported quality e4 Gonella et al. Vol. 00 No. 00 xxx 2023

and duration of end-of-life conversations). S.G. and Y.B. independently extracted all data, and solved disagreements by consensus with P.DG.

Primary Outcomes

The primary outcomes were process indicators of end-of-life conversations: occurrence, quality, timing, frequency or duration. Occurrence: the proportion of patients reporting, or with documented conversations. Quality: the quality of conversations either perceived by patients, family caregivers, or professionals, or according to their thematic content. Timing: time before death. Frequency: number of conversations per patient irrespective of the timing. Duration: length of conversations. Occurrence was expressed as percentage, quality as mean (SD) for quantitative measures (e. g., quality of care item) or thematic domains of conversations (e.g., values and goals, prognosis) for qualitative measures, timing as mean or median, frequency and duration as mean.

Data Analysis and Synthesis

Interventions were categorized into those targeting patients, professionals, or multiple stakeholders, and according to system level strategies e.g., alerts, electronic documentation template, and the number of sessions (one-time vs. multiple-sessions interventions).

We used prevalence ratios for occurrence, standardized mean difference (SMD) for patient-reported quality and ratio of means (ROM) for duration of end-of-life conversations. A random-effect model meta-analysis with unrestricted maximum likelihood using the sample size as a weighting factor was performed when any of the primary quantitative outcomes was assessed in at least four studies. Hartung-Knapp method was applied. Heterogeneity was assessed using the Cochran Q test via a Mantel-Haenszel test and quantified by I² statistic. Subgroup analyses with random-effects models investigated the association between documented occurrence and persons targeted, system strategies, and number of sessions. Sensitivity analyses for outliers were also performed.

Funnel plots were used for publication bias. Results were considered statistically significant at 2-tail P < 0.05. R v 4.1.2 was used for all analyses.

Results

Review Process

A total of 4,663 articles were identified; 32 full texts remained after screening titles and abstracts. Seventeen were excluded (Table A2). Three additional articles were identified from reference lists; thus 18 studies were included (Fig. 1). These assessed five process

outcomes: occurrence, \$\frac{31,49-58}{31,49-58}\$ quality, \$\frac{49,50,52,53,55,56,58-63}{49,55,56,58-63}\$ timing, \$\frac{31,55,58}{31,55,58}\$ frequency, \$\frac{31,55}{31,55}\$ and duration \$\frac{62-65}{62}\$ of end-of-life conversations (Table 1 and 2). Two articles could not be included in any meta-analyses: one \$\frac{60}{62}\$ did not quantify the quality of conversations, and one \$\frac{61}{62}\$ did not provide overall estimates of changes in the patient-reported quality of conversations.

Characteristics of Included Studies

Thirteen $^{31,49-60}$ studies were conducted in the United States, two 62,63 in Australia, one 64 in the Netherlands, one 65 in France, and one 61 in Japan; all but two 51,57 were published after 2005. Ten $^{49-52,55,59,60,63}$ studies were classified as high, five 31,53,56,57,62 as moderate and three 54,58,61 as low quality (Table A3). Thirteen $^{31,49-52,54-57,61,62,64,65}$ studies were hospital-

Thirteen^{31,49–52,54–57,61,62,64,65} studies were hospital-based, three^{59,60,63} involved both clinics and inpatient services, and two^{53,58} only clinics.

Studies involved patients with cancer, ^{50-52,54-57,59} -65 end-stage pulmonary, ^{31,49,51,52,56,57,59,65} cardiac, ⁵¹ -53,56,57,59,65</sup> liver, ^{51,52,59,65} renal, ^{52,56,57} or metabolic disease, ^{52,57} or multiple severe comorbidities ^{51,58,59}. Only one study involved patients with dementia, ⁵⁶ gastrointestinal, ⁵⁷ infectious, ⁵⁷ neurologic, ⁵⁷ or rheumatologic ⁵⁷ end-stage disease. Among professionals, studies most frequently involved oncologists, ^{50-52,55,60,62-64} or internists ^{49,51,52,57,59} (Table 1, A4).

Definition of End-of-Life Conversations

In the included studies, end-of-life conversations were defined as discussions, conversations, communications, talks, or information about symptom management, because $5^{1,53}$ treatment plans, because $5^{1,53,61}$ end-of-life care, $5^{7,59,60,63}$ goals of care, because $5^{1,53,61}$ end-of-life care, because $5^{1,53,61}$ goals of care, $5^{1,53,61,62,64}$ treatment options with their benefits and harms, $5^{1,55,61,62,64}$ illness trajectory and prognosis, $5^{1,53,61,62,64}$ and patients values or care preferences $5^{1,49-51,55,56,58,62,64,65}$ (Table 1).

Interventions to Promote End-of-Life Conversations

In all, 12 unique interventions across 18 studies were identified. The interventions targeted multiple stakeholders (n = 10), $^{31,49,52,53,55,60,62-65}$ professionals only (n = 7), $^{50,51,54,56-59}$ or patients only 61 . Four studies 60,62,63,65 involved family caregivers.

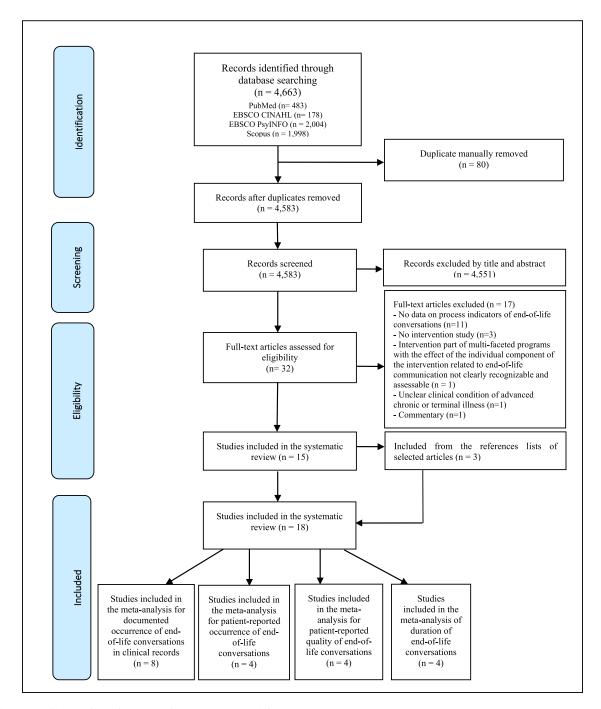


Fig. 1. PRISMA flow-chart depicting the main stages of the systematic review process. Page MJ, McKenzie JE, Bossuyt PM. et al. The PRISMA 2020 statement: An updated guideline for reporting systematic reviews.

individual- or small group-based reflective discussions $(n=5)^{54,57,59,60,64}$ provision of a prognostic estimate $(n=1),^{51}$ or palliative care consultation $(n=1)^{54}$ targeted only professionals. Five $^{51,55-58}$ studies also introduced strategies at the system level. System strategies included email remainders, alerts, or administrative prompts $(n=2),^{56,57}$ an electronic documentation template for conversations $(n=2),^{55,58}$ and a trained nurse who facilitated conversations $(n=1)^{51,58}$. Eight 31,49,52,53,57,61,63,65 studies assessed one-time

interventions while ten^{50,51,54–56,58–60,62,64} multiple-session interventions. Interventions were delivered by nurses only,^{51,53,62} physicians only,^{50,63,65} experienced trainers (undefined qualification),^{56,64} or multi-professional teams^{43,54,55,57,59,60} (Table 1, 3, and A5).

Outcomes of Interventions Aimed at Promoting End-of-Life Conversations

All funnel plots exhibit asymmetry, as depicted in Figure A1. While asymmetry has traditionally been

$Table \ 1$ Characteristics of Included Studies

Author(s)				Study Characteris	stics					Study Samp	ole Characterist	ics		Main Findings
(Country, year) Quality Score* Applicability**	Design / Study Period	Definition of EoL Conversation	Intervention	Who Delivered the Intervention	Process Indicator(s) of EoL Conversations Investigated	Data Source (Time of Data Collection)		Patients Underlying Disease	Life Expectancy (m)	Family Mem (N) Males (%) Age, years	(N) Males (%) Age, Years	Professional Professional Profile	Working Experience, Years ^a	
Au et al. (USA, 2012) High Mostly pragmatic	Cluster RCT / January 2004 to November 2007	Discussions about preference for EoL care	patient's survey about preference for EoL care was provided to physicians (n = 42) and their patients (n = 194) C: no patient-specific feedback for neither physicians (n = 50) nor	NA	Patient-Reported Quality of EoL Conversations Proportion of patients reporting occurrence of conversation about EoL preferences between patients and their physician or their surrogate	Patients' questionnaire (baseline and two weeks after the clinic visit)	Two hospitals / Not reported	N = 376 Chronic obstructive pulmonary disease	Not reported	-	N = 92 I: 42 M: 21 (50) Age: Not reported C: 50 M: 22 (44) Age: Not reported	Internists, lung specialists, geriatricians	•	1) Improved Quality of EoL conversations Assessed with the QoC Questionnaire 0–100 in the Intervention Group (Adjusted difference 5.74 points p = 0.03, Cohen effect size = 0.21); 2) More intervention patients reporting EoL conversations with their clinicians (Adjusted difference 27.4%, P<0.001); 3) More intervention patients reporting EoL conversations with their surrogate (Adjusted difference 10.9%, p<0.01).
Bickell et al. (USA, 2020) High Mostly pragmatic	Cluster RCT / Not reported	Conversations on eliciting patient values about care and treatments	patients (n = 182) I: physicians received a 2-hour small group role play session+4 coaching visits (n = 11) C: no training (n = 11) All intervention and control physicians attended a didactic lecture about the importance of conducting EoL discussions	,	Proportion of patients reporting occurrence of EoL conversations Proportion of patients reporting high-quality EoL conversations	Patients' questionnaire (within days after the clinic visit and at 6 months)	Three hospitals / Not reported	N = 265 Advanced cancer	< 24		N = 22 I = 11 M: 7 (64) Age: 44.3 (8.9) C = 11 M: 8 (73) Age: 43.5 (10.7)	Oncologists	I: 17.3 (Not reported) C: 17.0 (Not reported)	1) No difference in the occurrence of EoL conversations in intervention patients compared to control patients (52% vs. 48% OR 0.98, 95% CI, 0.95 – 1.01); 2) No difference in the occurrence of high-quality EoL conversations (scores of 9–10 on NRS 0–10) in intervention patients compared to control patients (62% vs. 67%, P=0.37) 3) No difference in the composite score of occurrence or quality or EoL conversations in intervention patients (Adjusted OI 0.84, 95% CI 0.57–1.23); 4) At 6 months, no difference in the mean number of emergency department visits and hospitalization, median time between hospice enrollment and death, chemotherapy administration, and intensive care unit admission.

 $Interventions\ to\ Promote\ End-of-Life\ Conversations\ in\ Clinical\ Encounters$

Table 1 Continued

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Aunot (s) (Country, year) Quality Score* Applicability***	Design / Study Period	Design / Study Definition of Intervention Period EoL. Conversation		Who Delivered the Intervention	thy cina accessings Who Delivered Process Indicator(s) Data Source (Time the of EoL of Data Collection) Intervention Conversations Investigated	Data Source (Time of Data Collection)	Setting/ Size (N)	Patients Underlying Disease	Life Expectancy (m)	H Y	ers (N) Males (%) Age, Years	Professionals Professional Profile	Working Experience, Years ^a	Stunner i men
Clayton et al., (Australia, 2007) High Mostly pragmatic	Parallel-group RCT / October 2002 to August 2004	Discussions about prognosis and EoL issues	I: patients (and their family caregiver if present) were given a question prompt list 20–30 minutes before their consultation with the physicians were asked to actively endorse and refer to the question prompt list using a standardized consultation (n = 92)	Physician	Count of overall questions, or paliative care service, physical symptoms, treatment, prognosis, quality of file, support, concerns about professional care, concerns about professional care, concerns and professional care, their family caregivers and for patients and for patients and for patients and for patients and cora finally caregivers. Total number of items discussed buration of consultation	Audio-recorded	Nine palliative care services (mostly outpatient clinics) / Not reported	N = 174 Advanced cancer	Not reported	N = 123 I: 70 M: NR Age: NR C: 53 M: NR Age: NR	N = 15 M: Not reported Age: Not reported	Palliative care physicians	Not reported	1) More overall questions (P < 0.0001), questions about palliainte care service (P < 0.0001), prognosis (P = 0.004), quality of life (P < 0.0001), and support (P < 0.0001), and intervention patients: 2) More overall questions about palliative care service (P = 0.03), prognosis (P = 0.03), questions about palliative care service (P = 0.03), prognosis (P = 0.05), and caregiver issues (P < 0.0001) in intervention family caregivers; 3) Overall, 22% more items discussed during consultations with intervention patients compared to control (P < 0.0001); 4) Increased consultation duration (37.8 minutes in the control group, s. 30.5 minutes in the control group, ratio 1.24 45% CI 1.09-1.41); 5) Not difference in patients anxiety, their satisfaction with communication and overall
Connors et al., (USA, 1995) High Mostly Pragmatic	Cluster RCT / January 1992 to January 1994	Discussions to elicit preferences, improve under standing of outcomes, en courage attention to pain control, and facilitate planning for future decisions	I: physicians received prognostic information and patient preferences for EoL care. A skilled nurse had multiple contacts with the patient, family, physician and bospital staff to elicit and document perferences, improve understanding of understanding of encourage attention to pain control, and facilitate planning for future decisions C. usual care	Nurse	Proportion of patients patients reporting occurrence of EoL conver sations	Patients interview or Five hospitals / N = 4804 family caregivers Not Oncol interview when reported pulmor patient interview was not possible fiver est does and 7 and again between days 6 coma, and 7 and again coma, and 15 after ongan enrollment) system failure sepsis	Five hospitals / Not reported	N = 4804 Oncologic, pulmonany, cardiac and liver end- stage disease, coma, or multiple organ system system stallure with sepsis	Not reported		N = 27 phy sécian's secian's groups I: 16 physician specialty groups C: 11 physician specialty groups groups groups	Internal medicine, pulmono logy/ medical medical care unit, oncology, surgery, and cardiology	Not reported	measures of achievement of information needs at 24 hours and 3 weeks. 1) No difference in the prevalence or timing of decumented DNR orders (Adjusted ratio of median time 1.02, 95% CJ 0.90–1.15); 1) No difference in the proportion of intervention patients of ramily caregivers reporting discussed cardiopulmonary resuscitation preferences compared to control patients (40% as 37%); 2) No difference in the median days spent in innersive care units, comatose, or receiving mechanical ventilation, neither in moderate to severe self-reported pain during the five-yrs study period.

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Table 1 Continued

Main Findings		1) Comparing intervention to control, no difference in reported quality of EOL conversations on the QoC questionnaire score (0-10) for patient (Adjusted b 0.38, 45% CI -0.77 to 0.98, P= 0.81); and chiridan (Adjusted b 0.11, 95% CI -0.77 to 0.98, P= 0.81); and chiridan (Adjusted b 0.11, 95% CI -0.78 to 0.98, P= 0.81); and chiridan (Adjusted b 0.19, 95% CI -0.08 to 0.47, P= 0.17); 2) No association between the intervention and patients' physical and mental status, or family careguers' depression score; increased depression score; increased depression among nitervention patients	1) Wore intervention patients reporting occurrence of Ed. conversations (7.4% vs. 31%, Adjusted #1.25 [19% CI 0.34-1.156], P< 0.001). 2) More intervention patients with documented Ed. conversations (62% vs. 17%, Adjusted #1.25 [195% CI 0.92-1.58], P< 0.001); (95% CI 0.92-1.58], P< 0.001); (195% CI 0.92-1.58], P< 0.001); conversations assessed with the Coloquestionmaire (a priori selection of 4 tiems rated from 0 to 10) in intervention patients (4.59 vs. 2.13, Adjusted #5.02 [195% CI 0.488.37], P=0.01); 4) No difference in depression or	anxiety at 5 and 6 monus.
Ma	Working Experience, Yeans ^a	I: median 1.0 1) IQR (1.2) C: median I: 0 IQR (1.2) 2) 2) 2)	Not reported 1)	
ş	Professionals Professional Profile	Internal medicine residents and nurse practitioner trainees	Family medicine, internal medicine, oncology, pulmo nology, cardiology, cardiology, represented particles, rephrology, geriatrics	
Study Sample Characteristics) Males)) Age, ars	N = 445 E 211 M: 91 Age: 30.5 (5.8) C; 234 M: 98 (42) Age: 30.3 (4.8) Climicans evalua roos N = 890	N = 124 M: 58 (46.5) Age: 47.2 (9.6) I = 59 M: Not reported Age: Not reported Age: Not reported Age: Not reported Age: Not reported	
Study Sampl	Family Members (N) Males (N) (%) Age, (%) years Ye	N = 898 E: 412 M: 106 (26) Age: 56.8 (13.7) C: 486 M: 124 (26) Age: 56.6 (13.5)		reported
	Life Expectancy (m)	12 to 24	<294	N = 80 Heart failure
	Patients Underlying Disease	N = 1717 Oncologic, pulmonary, cardiac or liver end- stage disease, or multiple severe comorbi dities	N = 494 Oncologic, pulmonary, cardiac, liver renal, metabolic end-stage disease	One heart failure compatient clinic/600 patients each year
	Setting/ Size (N)	Primary care clinics and inpatient services / Not reported	Seven hospitals, one cancer center, two outpatient network / Not reported	question question naire and electronic medical record review (baseline and two weeks after the clinic visit)
	Data Source (Time of Data Collection)	Patients, family caregivers and clinicians surveys clinicians surveys furning the 6-munith period preceding intervention and in the 10 months following the intervention)	Patiens' questionnaire questionnaire (two weeks after the clinic visit) Electronic medical record review (from the dinic visit months) months)	Proportion of patients with documented occurrence of EoL romersations Patient-reported quality of EoL conversation s
stics	Who Delivered Process Indicator(s) the of EoL Intervention Conversations Investigated	Patient-reported quality of EoL conversations Family caregiver-reported quality of EoL conversations Clinician-reported quality of EoL conversations	Proportion of patients reporting occurrence of EoL Conversations Proportion patients with documented occurrence of EoL conversations Patient-reported quality of EoL conversation s	Миже
Study Characteristics	Who Delivered the Intervention	Physician Nurse	¥.	1: patients received nurse-led telephone-based previsit coath ing-tone pag patient activation outline constructed during the call and shared with both the patient and their physician physician
	Intervention	I: physician and nurse trainees received eight 4-hour sessions including overview didactic with demonstration role-play, simulation and reflective discussions (n = 211) (c usual education (n = 234)	I. a patient-specific l-page communication tips based on a previous survey was sent to patients (n = 249) and dinicians (n = 65) one week and 10°2 working days prior to the clinic visit, respectively (c. patients completed the same survey but no information was provided to patients (n = 288) or the survey but the same survey but no information was provided to patients (n = 288) or the same survey but th	cunctant (H = 0.) management, and advance care planning
		Discussions about palliative and EoL care	Talks with patients about their prognosis or goals of care	
	Design / Study Definition of Period Conversation	Cluster RCT / October 2007 to January 2013	Chuster RCT / February Pol14 to May 2016	Discussions about illness trajectory, prognostic estimates, symptom
Author(s)	(Country, year) Quality Score* Applicability***	Curtis et al. (USA, 2013) High Mostly explanatory	Gurtis et al. (USA, 2018) High Mostly explanatory	Parallel-group RCT / twoyrs period

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Table 1 Continued

Author(s)		Study Characteristics	stics					Study Sample Characteristics	Characteristi	CS		Main Findings
/ear) ore* ty**	Design / Study Definition of Intervention Period Eo.L Conversation	Who Delivered the Intervention	Process Indicator(s) of EoL Conversations Investigated	Who Delivered Process Indicator(s) Data Source (Time the of Fol. of Data Collection) Intervention Conversations Investigated	Setting/ Size (N)	Patients Underlying Disease	Family Me Life (N) Males Expectancy (%) Age, (m) years	Family Members (N) Males (N) (%) Age, (%) years Yea	ers (N) Males (%) Age, Years	Professionals Professional Profile	Working Experience, Years ^a	,
		along with suggestions for										
		addressing barriers +										
		patient activation,										
		skills enhance										
		ment, and										
		conversa										
		tion										
		(n = 41)										
		C: regular visit (n = 39)										
												1) More intervention patients with
												documented EoL conversations (73% vs. 2.6% , $P < 0.001$);
												2) Higher quality of EoL
												Conversations assessed with the
												intervention patients ($P = 0.03$,
												Cohen effect size = 0.43);
												general conversations between
												intervention and control
												patients ($P = 0.76$, Cohen effect
												4) No difference in completion of
												advance directives between
												intervention and control
												patients (16.1% vs. 7.7%, $P = 0.24$);
												5) No difference in palliative care

SI		1) Improved patient-centered communication in the intervention group (Aditisted b intervention group (Aditisted b 0.03, 1034 [95% CI 0.06–0.02], P = 0.02) corresponding to 5.7 additional "engaging" statements (F44%), 0.6 statements (F44%), 0.6 additional responses to emotion (+71%), and 1.4 additional asturents regarding prognosis and treatment choices (+88%); 2) No difference in quality of life and aggressive care.	
Main Findings		I) Improve communities of mineryear intervent of the control of co	
	Working Experience, Years ⁴	Not reported	
iis	Professionals Professional Profile	Oncologists	
Study Sample Characteristics	bers (N) Males (%) Age, Years	N = 38 I = 19 M: 12 (6.3) Age: 43.9 (10.3) C = 19 M: 15 (79) Age: 45.3 (9.8)	reported
StudySamr	F O O V	N = 194 M: Not reported Age: Not reported	Advanced Not
	Life Expectancy (m)	<u>~</u>	N = 172
	Patients Underlying Disease	N = 265 Advanced cancer	Medical oncology unit of a 804-bod university hospital
Conunuea	Setting/ Size (N)	Community- based cancer clinics, academic nedical conterns and community hospital Not reported	Medical record Medical review / Not oncol reported unito 804-bh univer hospin
3	Data Source (Time of Data Collection)	Audio-recorded consultation (after the first training session)	Proportion of patients with decumented occurrence of EoL.
tics	Process Indicator(s) Data Source (Time of Eo.L. of Data Collection) Conversations Investigated	Composite measure of quality communication (z scores of engaging patients in consultation [Active Patient Participation [Active Patient Participation [Active Patient Participation (Coding). responding to patients [Verona VR-CoDE3], informing patients about prognosis and treatment choices [Prociees [Procie	Nursee () Physician Social worker
Study Characteristics	Who Delivered the Intervention	worker	I: residents, medical students, murse practitioners, and physicians assiants received monthly 45-mining didactic regions skills training didactic regroup for 10 months remonthly for 10 months palliative care consultation for 8 months months
S	Intervention	I: oncologists received a two-sessions in- office training (1.75 video and feedback from standardized patients portraying roles of patients +focused learner- centered reflective feedback Patients and their caregivers received a single one-boar coaching seesion incorporating a question prompt list +up to 3 follow-up phone calls in the next 3 months. C: no training	Communication between the patient or patient surrogate and a treating physician or advanced practice provider, with content covering some of the flowing treatment goals and values, prognosis, cancer treatment choice, life sustaining treatment choice, life sustaining treatment choice, ire sustaining treatment choice, or hospice or confort care options.
	Definition of EoL Conversation	Discussions about disease course, prognosis, treatment decisions and EoL. care	
	Design / Study Definition of Period EoL. Conversation	Cluster RCT / August 2012 to October 2015	Pre-post study / July 2015 to May 2016
Author(s)	(County, year) Quality Score* Applicability**	Epstein et al. (USA, 2017) High Mostly explanatory	Hanson et al. (USA, 2017) Low Mostly pragmatic

 $Interventions\ to\ Promote\ End-of-Life\ Conversations\ in\ Clinical\ Encounters$

Table 1 Continued

Who Delivered Process Indicator(s) Da the of EoL of Intervention Conversations Investigated	Study Sample Characteristics
	Process Indicator(s) Data Source (Time Setting/ Size Underlying Life (N) Males (N) Males Professionals Working of EoL Conversations (N) Disease Expectancy (%) Age, (%) Age, Profile Experience, (m) years Years Years
Physicians' 10-hour Experienced Duration of training consisted trainer consultation group sessions (ridecessrole play), and the personal feedback on a videotaped consultation room on a videotaped consultation room consultation and a consultation and a consultation and a consultation and a consultation and experience described by the physician training training the physician training train	f Videonped Seven hospitals N = 194 < 12 . N = 31 Oncologiss 7.7 (9.0) Tribution / Not Advanced Age: 41.5 (9.5) Age: 1.15 M: Not reported Age: Not reporte

$Table\ I$

Author(s)				Study Characteristics	tics					Study Sampl	Study Sample Characteristics	cs		Main Findings
(Country, year) Quality Score* Applicability***	Design / Study Period	Design / Study Definition of Period EoL Conversation	Intervention	Who Delivered the Intervention	Process Indicator(s) Data Source (Time of EoL Conversations Investigated	Data Source (Time of Data Collection)	Setting/Size (N)	Patients Underlying Disease	Life Expectancy (m)	Family Members (N) Males (N) (%) Age, (%) years Ye	ers (N) Males (%) Age, Years	Professionals Professional Profile	Working Experience, Years ^a	
Lakin et al. (USA, 2017) Low Mostly pragmatic	Controlled dinical trial / January 2014 to May 2015	Discussions about prognosis, values, goals, or car preferences	1: clinicians received 2.5 hours role playbased training on the use of a conversation guide +monthly coaching (via telephone, email or in-person) +system changes (i.e., surprise question, electronic documentation template, nurse coordinator was notified about played about about played about	Palliative care experts	Proportion of patients with documented occurrence of at least one EoL conversation before death Timing of the first documented EoL conversations before death Comprehen siveness of EoL conversations Proportion of patients with documented occurrence of EoL conversations in the advance care planning module of the electronic medical record	Electronic medical record review (dead patients)	Fourteen primary care chinics / Not reported reported	N= 178 Multiple sewere comorbi dities	45		1: 59 M: Not reported Age: Not reported Age: Not reported Age: Not reported Age: Not	Physicians, nurse care coordina tors, social workers	Not reported	1) More intervention patients with documented occurrence of at least one conversation before death (62.4% vs. 42.9%, P = 0.00%; 2) No significant difference in the timing of the first documented conversation in intervention patients compared to control patients compared to control patients compared to control patients compared to control death vs. 129.2 d. respectively, 9 More comperhensive conversations about values and goals in intervention patients (mean 1.8 elements/patient vs. mean 1.8 elements/patient values and discussion of prognosis, life sustaining treatments, or Eo. More intervention patients with documented conversation in the advance care planning module of the electronic medical record (44.4% vs. 3%, P e 0.001);
Lautrette et al. (France, 2007) High Mostly pragmatic	Parallelgroup RCT / May 2005 to October 2005	Discussions about patients' wishes wishes aimed to promote under standing about the goals of care support	I: clinicians were asked Physicians to follow detailed guidelines for EoL confe errores; family rences; family rences; family rences; and a bereave ment information leaflet (n = 63) C: usual practice (n = 63)	Physicians	Duration of consultation	Clocking of family conference	22 Intensive Care (medical and surgical) Unix, 15 in teaching hospitals A in median mumber of bed (IQR)	N = 126 Gronic observe ive pulmo nary disease, chronic heart failure, cancer, cirrhosis	A few days	N = 126 1: 63 M: 12 (23) Age: median 54 (1QR 46-64) C: 63 M: 17 (30) Age: median 54 (1QR 47-58)	N = Not reported M: Not reported Age: Not reported	Not reported	Not reported	1) Longer family conference in the intervention group than in court of (median 30 [10R 19 –45] minutes vs. 20 [10R 15 –45] minutes vs. 20 [10R
Pre-post study / April 2004 to March 2008	Information including disease conditions, therapeutic measures, care procedures, and treatment plans	I: disclosure of Not reported more detailed information about cancer according to the baseline level of awareness:	Not reported	Patient- reported quality of communi cation between patient and family Patient- reported quality of	communication between medical professionals Patient reported quality of communication between patient/ family and medical professional	Patients' questionnaire (before and one week after disclosing information)	Two hospitals / Not reported	N = 73 Advanced cancer	reported	1				1) No improvement in the quality of any communication for disclosed group A patients on STAS-J (0-4). 2) Improved communication quality between patients and families, between medical professionals, and between patient/families and medical professionals for both disclosed group B and C patients on STAS-

 $Interventions\ to\ Promote\ End-of-Life\ Conversations\ in\ Clinical\ Encounters$

Author(s)				Study Characteristics	tics					Study Sampl	Study Sample Characteristics	ics		Main Findings	
(Country, year) Quality Score* Applicability***	Design / Study Definition of Period EoL Conversation		Intervention	Who Delivered the Intervention	Who Delivered Process Indicator(s) Data Source (Time of EoL of Data Collection) Intervention Conversations Investigated	Data Source (Time of Data Collection)	Setting/ Size (N)	Patients Underlying Disease	Life Expectancy (m)	Family Members (N) Males (N) (%) Age, (%) years Ye;	oers (N) Males (%) Age, Years	Professionals Professional Profile	Working Experience, Years ^a		
		Group A. disclosure disclosure disclosure disclosure (n = 8) Group B. disclosure of life-threatening disclosure of life-threatening disclosure of disclosure of prognosis (n = 40)												J (0-4) (P= 0.0085, P= 0.0062, P= 0.0013 and P= 0.02, P= 0.0082, and P= 0.027, P= 0.0082, and P= 0.0057, respectively); 3) No improvement in the quality in any communication in not disclosed group A and B patients on STAS4 (0-4); 4) Improved communication quality between medical professionals and between patients/families and medical professionals from the patients on STAS4 (0-4); 4) Group C patients on STAS4 (0-4);	= 0.0062, 02, 0057, 0057, the quality in not- id B patients cation da B patients tween d medical d medical andisclosed STASJ (0
Paladino et al. (USA, 2019) High Mostly pragmatic	Cluster RCI / 1 September 2012 to June 2016	about life- protonging protonging protonging protonging proton and proton and proton and preferences preferences	1: physicians received structured communication communication guide-2.5 hours skilled-based training program +coaching-system changes (i.e., surprise question, eerati remiders, electronic documentation, template) (in = 48); patients received no patients received no intervention on intervention of patients using the surprise question (in = 43); the patients (in = 85)	Palliative care faculty	Proportion of patients with documented occurrence of at least one EoL occursation before death Timing of the first documented EoL conversation before death Comprehen Compressions Frequency of documented EoL conversations in patients with documented occurrence of EoL conversations in an accessible module of the electronic medical record	Electronic medical record review (after death)	Three oncologic centres/ Not reported	N = 161 Advanced cancer	42>		N = 91 M: Not reported Age: Not reported G: 43 M: Not reported C: 43 M: Not reported Age: Not reported Age: Not reported Age: Not	Oncologists	mean 11 5 (95% CI 9.2- 13.8)	1) More intervention patients with documented occurrence of at least one conversation before least one conversation before death (96% vs. 79%, P = 0.005); 2) Earlier conversations in the intervention group than in the control group (median 143 [1Qk, 71–829] d before death vs. 71 [1Qk, 83–166] d, respectively, P < 0.001); 3) More intervention patients with documented conversation about values goals (89% vs. 44%, P < 0.001), prognosis (1lle-sustaining treatment) preferences (63% vs. 0.001), and file-sustaining treatment preferences (63% vs. 92%, P = 0.004); no difference in doctournented conversation about documented conversation about Eol. ACP (80% intervention so 68% control, P = 0.08); 4) Higher comprehensiveness (P < 0.001) and firequency (mean 3.1 [95% CI 2.5.46] conversation per patient vs. mean 2.1 [95% CI 1.4.28] conversation in the accessible module of the electronic medical record (61% vs. 11%, P < 0.001).	an un before a nu before of at an un before of at an un before be a loud by a sin the han in the han in the han in the large of the lar

Gonella et al.

Table 1
Continued

Main Findings	Professionals Professional Working Profile Experience, Years*	Not reported 1) No difference in documented conversations between intervention and control (OR 0.97, 95% CI 0.64-1.46, P= 0.8616); 2) Increased percentage of time among trained physicians to address emotions (from 50% to address from 42% to 55%), elicit understanding of illness (from 42% to 50%), and make recommendations (from 42% to 50%), and make recommendations (from 42% to 50%), and make recommendations (from 43% to 50%). 3) No difference in documented ACP in intervention compared to control (4.5% vs. 3.8%, OR 2.01, 95% CI 0.96-4.21, P= 0.062); 4) Fewer overall (P<0.001) and 3.40, in intervention compared to control; no significant difference in length of stay, realizable of stay,	pominary care relation and hospice referral. 1) No difference in the proportion of patients with documented cocurrence of at least one EoL conversation before death between intervention and control (75% ss. 72%. Adjusted OR = 1.30, 95% CI 0.58 – 2.92, P = 0.53); 2) No difference in the odds of completing an advance directive (OR 1.30, 95% CI 0.614 – 2.77); 3) Documentation of 451 unique EoL conversations for 115 of 157 (75%) patients; 4) No difference in the mean frequency of documented EoL conversations in intervention patients compared to control patients (CRS) 3.2), P = 0.47);
Study Sample Characteristics	Eamily Members Professional Life (N) Males (N) Males Professional Expectancy (%) Age, (%) Age, Profile (m) years Years	Not	reported reported
	Patients Data Source (Time Setting/ Size Underlying of Data Collection) (N) Disease	Medical record One hospital / N=428 review / Not Not Metastatic reported cancer, dementia, congestive heart failure, chronic c	Medical record One hospial / N = 157 review (dead Not Ghronic patients) reported obstructive pulmonary disease
Study Characteristics	Who Delivered Process Indicator(s) Da the of EoL Intervention Conversations investigated	Communica Proportion of M tion coach patients with documented occurrence of EoL Comversations Compersions conversation s conversation s	NA Proportion of MA patients with documented documented occurrence of at least one EoL conversation before death Prequency of documented EoL conversation in Timing of documented EoL conversation is before death
	Design / Study Definition of Intervention Period EoL Conversation	Chuster RCT / Conversations I: physicians received July 2017 to about goals, alerts and 2-hour March 2018 preference communication and coaching provided prognosis in 3 sessions (n = 8) that affect C: no training (n = 7) restment decisions	randy of a help patient-specific cluster RGT patients and feedback form a surrogate based on a previous 2004 to decision patient's survey September makers prepare to for EoL care was make in-the provided to moment to physicians and their decisions patients based on Cro to patients and their decisions patients and their sylvatic one's values feedback for and makers and makers patients physicians and their patients for care in or patients for care
Author(s)	(Country, year) Quality Score* Applicability***	Pollak et al. (USA, 2019) Moderate Mostly pragmatic	Reinke et al. (USA, 2017) Moderate Mostly pragmatic

Table 1 Continued

Author(s)				Study Characteri	stics					Study Samp	le Characterist	ics		Main Findings
(Country, year) Quality Score* Applicability**	Design / Study Period	Definition of EoL Conversation	Intervention	Who Delivered the Intervention	Process Indicator(s) of EoL Conversations Investigated	Data Source (Time of Data Collection)	0.	Patients Underlying Disease	Life Expectancy (m)	Family Mem (N) Males (%) Age, years	bers (N) Males (%) Age, Years	Professional Profile	Working Experience, Years ^a	
Shorr et al. (USA, 2000) Moderate Mostly pragmatic	Pre-post study / pre- intervention cohort February to May 1995; post- intervention cohort April to June 1997	Discussions regarding EoL care and EoL issues	I: Physicians received administrative prompts to encourage EoL conversations and had a mandatory one-day educational seminar on EoL issues based on didactic lectures, small group discussions and role-playing	Ethics committee members	Proportion of patients with documented occurrence of EoL. conversations	Medical record review / Not reported	One hospital / Not reported	N = 305 Cardiac, pulmonary, gastrointes tional, infectious, renal, oncologic, neurologic, rheumato logic, or metabolic end-stage disease	<36	-	N = 115/167 (68.9%) complet ed the training M: Not reported Age: Not reported	Internists	Not reported	1) No difference in documented occurrence of EoL conversations in the pre-intervention cohort compared to the post-intervention cohort (34.8%, 95% CI 28.0–42.2 vs. 33.9%, 95% CI 25.7–43.1); 2) No difference in documented DNR orders in the pre-intervention cohort compared to the post-intervention cohort (28.8%, 95% CI 22.5-36.0 vs. 27.3%, 95% CI 19.8-36.3, P = 0.71).
Walczak et al. (Australia, 2017) Moderate Mostly pragmatic	Parallel-group RCT / Not reported	Discussions about prognosis and end-of- life care preference and future care options	I: Patients (n = 61) received two sessions of a nurse- led communication support programme (45- min face-to-face meeting based on a question prompt list and DVD discussing ACP+15 min telephone booster session) and caregivers could join; physicians were cued to use the question prompt list and question asking during consultation C: No communication support programme for patients (n = 49) nor physicians were cued to use the question prompt list and question asking	Nurse	Count of cues and questions for prognosis, EoL care future care options and overall for patients and their caregivers during consultations Duration of consultation	Audio-recorded consultations	Six cancer centres / Not reported	N = 110 Advanced cancer	2-12	M: Not reported Age: Not reported	N = Not reported M: Not reported Age: Not reported	Oncologists	Not reported	 More overall cues and questions (P = 0.025), cues and questions about prognosis (P = 0.010), Eol. care (P = 0.001), and future care options (P = 0.028) in intervention patients; More cues and questions about prognosis in intervention family caregivers (P = 0.014); Unchanged consultation duration (20.6 minutes in the intervention group vs. 20.4 minutes in the control group, P = 0.307); At one month, no difference in the overall, physical, social, emotional, or functional well-being between intervention and control.

Note. Process outcomes include occurrence, quality, timing, frequency, or duration of end-of-life conversations.

The primary outcome(s) of each study is/are in bold.

Abbreviations. ACP, Advance Care Planning; C, Control; CI, Confidence interval; d, day; DNR, Do Not Resuscitate; EoL, end of life; I, Intervention; IQR, Interquartile range; m, month; NA, Not applicable; NRS, Numeric Rating Scale; QoC, Quality of care; RCT, Randomized Controlled Trial; SE, Standard error; STAS-J, Japanese version of the Support Team Assessment Schedule.

^aExpressed as mean (SD) if not differently specified.

^{*}According to the Qualsyst critical appraisal criteria.

^{***}According to the PRagmatic Explanatory Continuum Indicator Summary (PRECIS) version 2 tool.

 ${\it Table~2}$ Process Outcomes of Interventions Aimed at Promoting End-of-Life Conversations between Professionals and Patients with Advanced Chronic or Terminal Illnesses or their Family Caregivers Assessed in the Included Studies

Author(s) (Country,			Pro	ocess Outcomes of	f Interventions Air	ned at Promoting E	nd-of-Life Conversatio	ons		
year)	Occur	rence of End-of-Life Conver	rsations		Quality of End	l-of-Life Conversatio	ons	Timing of End-of- Life Conversations	Frequency of End- of-Life Conversations	Duration of End-of-Life Conversations
	End-of-Life Conversations Documented in	Occurrence of End-of- Life Conversations Documented in a Specific Module of the Electronic Medical Record		Quality of End- of-Life Conversations According to Their Content	Reported Quality of End- of-Life	Reported Quality	Physician-Reported Quality of End-of- Life Conversations	Timing of End-of- Life Conversations	Frequency of End- of-Life Conversations	Duration of End-of-Life Conversations
Au et al. (USA, 2012) Bickell et al. (USA, 2020)			√ √		√ √					
Clayton et al. (Australia, 2007)			,	✓						✓
Connors et al. (USA, 1995)			√							
Curtis et al. (USA, 2013)					\checkmark	✓	\checkmark			
Curtis et al. (USA, 2018)	✓		✓		\checkmark					
Doorenbos et al. (USA, 2016)	✓			,	✓					
Epstein et al. (USA, 2017)				√						
Hanson et al. (USA, 2017)	√									
Henselmans et al. (The Netherlands, 2020)										√
Lakin et al. (USA, 2017) Lautrette et al. (France, 2007)		√		✓				√		
Nakajima et al. (Japan, 2015)					√					
Paladino et al. (USA, 2019)	✓	✓		\checkmark				\checkmark	\checkmark	
Pollak et al. (USA, 2019)	✓			✓						
Reinke et al. (USA, 2017)	✓							✓	✓	
Shorr et al. (USA, 2000) Walczak et al. (Australia, 2017)	✓			\checkmark						√

Interventions to Promote End-of-Life Conversations in Clinical Encounters

Table 3 Interventions Aimed at Promoting End-of-life Conversations Assessed in the Included Studies and Their Target

Author(s)	Target ing patient s	Targeting professionals	Targeting multiple stakeholders	Targeting the system	Multiple
(Country, year)	-		Patients/Family caregivers Professionals		-session interven tion vs
	Disclosure of more detailed information about clinical conditions	Patiens-specific information motor and set information preferences. Preferences of the proposals estimates pecific prognostic prognostic estimates. Roade pilos information in the prognostic prognostic estimates pecific prognostic estimates pecific prognostic estimates pecification estimates prognostic estimates and prognostic estimates and prognostic estimates e	information from the profession of the professio	Electronic documentation template Email reminders' alerts / administrative prompts Trained nurse facilitating end of-life conversations	one-time interven tion
Au et al. (USA, 2012)			~		One-time
Bickell et al. (USA, 2020)		√ √ √			Multiple- session
Clayton et al., (Australi a, 2007)*			✓		One≺ime
Connors et al., (USA, 1995)		✓ ✓		✓	Multiple- session
Curtis et al. (USA, 2013)		√ √ √			Multiple- session
Curtis et al. (USA, 2018)			✓ ✓		One-time
Dooren bos et al. (USA, 2016)			✓ ✓ ✓		One-time
Epstein et al. (USA, 2017)*			✓ ✓ ✓		Multiple- session
Hanson et al. (USA, 2017)		√ √ √			Multiple- session
Hensel mans et al. (The Netherla nds, 2020)			✓ ✓ ✓ ✓ ✓		Multiple- session
Lakin et al. (USA, 2017)		✓ ✓		√ ✓	Multiple- session
Lautrett e et al. (France, 2007)§			✓ ✓		One-time
Nakaji ma et al. (Japan, 2015)	√				One-time
Paladin o et al. (USA, 2019)			✓	✓	Multiple- session
Pollak et al. (USA, 2019)		✓		√	Multiple- session
Reinke et al. (USA, 2017)			✓		One-time
al. (USA, 2000)		√ √ √		✓	One-time
Walczak et al.			✓ ✓ ✓ ✓		Multiple- session
(Australia, 2017)*					

^{*} Family caregivers could join the programme together with the patient § Only family caregivers were involved

linked to publication bias and selective outcome reporting, it can also be caused by heterogeneity. Methodological differences can contribute to heterogeneity, with smaller studies, ⁵³ lower quality studies, ^{54,58} or studies with mostly explanatory design ^{52,53} tending to show larger effects. This is supported, in Figure A1, by studies reporting large effects also have larger standard errors.

Occurrence

Ten studies assessed documented occurrence of end-of-life conversations: eight $^{31,52-58}$ in clinical records, two 55,58 in a specific module of the electronic clinical record, and four $^{49-52}$ patient-reported occurrence. Seven of eleven $^{49,51-55,58}$ studies showed positive results, four of eleven 31,50,56,57 null results. Among the seven studies with positive results, one 53 had a wide CI with a small sample size and three 51,55,58 showed border-significant CIs.

Eight of eight^{31,52-58} studies contributed to the metaanalysis of documented occurrence in clinical records, with no significant effect of interventions (RR 1.54, 95% CI 0.84–2.84, I² 91%) (Fig. 2). Results were similar in sensitivity and subgroup analyses (Fig. A2).

Four of four $^{49-52}$ studies indicated no significant effect on patient-reported occurrence (RR 1.52, 95% CI 0.80–2.91, I^2 95%) (Fig. 2).

Quality

Patients', ^{49,50,52,53,59,61} family caregivers'⁵⁹ and professionals'⁵⁹ self-reports, and thematic content^{55,56,58,60,62,63} of end-of-life conversations were employed to assess quality. Quality was assessed using the Quality of Communication questionnaire, ^{49,52,53,59} the Support Team Assessment Schedule-Japanese

version, 61 the Numeric Rating Scale $0-10^{50}$ or a composite score of communication measures, 60 or coding the content of conversations in thematic domains. 55,56,58,62,63

Three of six studies 49,52,61 showed positive quantitative results, the other three of $\sin^{50,53,59}$ null results. Only $\cos^{49,52}$ studies with positive results contributed to the meta-analysis. In all, four of $\sin^{49,52,53,59}$ studies contributed to the meta-analysis, which showed no effect on patient-reported quality (SMD 0.83, 95% CI -1.06 to 2.71, I^2 99%) (Fig. 2).

Quantitative evidence showed that interventions targeting only professionals did not improve the quality of conversations perceived by family or professionals. ⁵⁹ In all, six of six 55,56,58,60,62,63 studies showed positive qualitative results based on the thematic domains of the conversations. Interventions targeting multiple stakeholders were more likely to increase the number of relevant domains (e.g., values/goals, life-sustaining treatment preferences) discussed^{55,63} and the time trained professionals addressed them,⁵⁶ compared to interventions targeting only professionals.⁵⁸ Also, interventions targeting multiple stakeholders increased the number of statements to inform patients about prognosis and treatment choices (+38%), engage patients in conversation (+44%), and respond to patients' emotions (+71%);⁶⁰ patients' and their family caregivers' cues and questions about end of life and end-of-life care also increased. 62,63

Timing

Three^{31,55,58} studies assessed timing. Conversations took place earlier when interventions targeted multiple stakeholders (median 143 days)⁵⁵ compared to involving only professionals (mean 133 days).⁵⁸

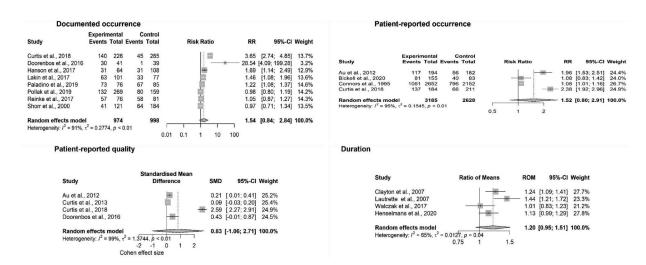


Fig. 2. Meta-analyses of interventions aimed at promoting documented occurrence in clinical records, patient-reported occurrence, patient-reported quality, and duration of end-of-life conversations. CI, Confidence interval; RR, Risk ratio; ROM, Ratio of means; SMD, Standardized mean difference.

Frequency

Of the two^{31,55} studies targeting multiple stakeholders, one⁵⁵ highlighted higher frequency of end-of-life conversations (mean 3.1 [95% CI 2.5–3.6] per intervention patient vs. 2.1 [95% CI 1.4–2.8] per control patient, P = 0.02), while the other³¹ found no difference (mean 3.1 (SD 4.0) vs. 2.6 (SD 3.2), P = 0.47).

Duration

Four studies targeting multiple stakeholders assessed the duration of end-of-life conversations. ^{62–65}

Two of four 63,65 studies showed positive results, the others 62,64 null results. Four of four 62,65 studies contributed to the meta-analysis, which showed no effect on duration (ROM 1.20, 95% CI 0.95–1.51, I² 65%) (Fig. 2).

Patients- and Family Caregivers-Related Care Outcomes

Thirteen of eighteen studies assessed the impact of interventions on outcomes ^{50,54,56,59,60,62–65} (Table A6).

Four^{54,56,59,65} studies (three^{54,56,59} targeting professionals only and one⁶⁵ multiple stakeholders) found significant changes with mixed results: decrease in overall (P < 0.001) and 30-day readmission rates (P = 0.009) with improvement of the content without changes in the occurrence of conversations;⁵⁶ increased occurrence of conversations in clinical records with increased palliative care consultation (P = 0.026) but reduced screening of constipation (P = 0.041);⁵⁴ decreased patients' post-traumatic stress disorder (P = 0.01), anxiety (P = 0.02), and depression (P = 0.003) with longer conversations;⁶⁵ increased patients' depression (P = 0.006) without improvement in patient, family caregivers, or clinician-reported reported quality of conversations.⁵⁹

Four^{31,51,53,57} studies assessed the impact of interventions on advance directives and one⁵⁶ on ACP. No difference in completion of advance directives neither in documented ACP was found.

Applicability of the Interventions

Fourteen 31,49–51,54–58,61–65 studies were mostly pragmatic, four 52,53,59,60 mostly explanatory. Among the domains of PRECIS-2 tool, eligibility criteria, recruitment modalities, flexibility in engagement, primary outcome, and primary analyses were mostly pragmatic. Setting, expertise and resources needed, flexibility in the delivery, and follow up were mostly explanatory (Table A7).

Discussion

Our systematic review on end-of-life conversations found interventions were mostly pragmatic with no effect on the occurrence, patient-reported quality or duration of conversations, regardless of who was targeted, the introduction of system strategies, or the number of sessions. The meta-analyses showed high heterogeneity likely due to wide variability in contexts, patients' clinical conditions, and interventions delivered. However, qualitative and quantitative evidence indicated that when interventions targeted multiple stakeholders, conversations were earlier and more comprehensive (e.g., number of relevant domains addressed or information about prognosis and treatment choices), regardless of the strategies employed; 55,60,62 unfortunately, these positive results could not be tested in meta-analyses to ascertain significant differences for the lack of power. Instead, the effect of interventions on frequency 31,55 was limited and conflicting.

Process indicators of occurrence, quality, timing, frequency, and duration provide clear and objective measures on how end-of-life conversations are delivered, and comply with the evaluation criteria of reliability, validity, responsiveness to change, and ease of implementation to effectively guide quality improvement efforts. ^{19,20}

Documented occurrence of conversations in clinical records was the most frequent process outcome, 31,52-58 followed by patient-reported quality, 49,50,52,53,59,61 and thematic content of conversations. 55,56,58,60,62,63 This may reflect recommendations in most international palliative care guidelines to clearly document conversations and their content.²⁵ Tracking what has been discussed, it is helpful to put at the front elements of shared decision-making such as the patient's understanding of illness and end-of-life care options, and their values and care preference.⁶⁶ Unfortunately, several domains of conversations relevant to direct end-of-life care are poorly discussed: only half of the conversations covers psychological problems and less than 40% addresses spiritual and existential issues. Also, options in terms of palliative care are not discussed at all in 45% of conversations. 6 Measuring the process rather than measuring only the final outcome -advance directives completionthat is often the result aimed for, allows to verify if the care goals are consistent with the patient's stated preferences. A multidisciplinary expert panel ranked consistency between documented care goals and the patient's stated preferences as the most important of a list of quality indicators for documentation of end-of-life communication and decision-making.¹³

Also, there is consensus that such information should be easily accessible by professionals within and beyond the current team involved in the care and cover conversations that have taken place at any time point over the disease trajectory. ⁶⁷ Care for adults nearing the end of life is indeed coordinated by health and social care professionals in various services and organizations that would benefit from knowing about any conversations about prognosis, treatment goals, and

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care plans.¹⁵ To support that patients' wishes are honored, it is crucial that the documentation is easily accessible.^{68,69} A structured, specific module on end-of-life conversations in the electronic clinical record may promote greater accessibility, and be used to monitor the effect of an intervention on process indicators of conversations, such as timing, frequency, and key content. Only two^{55,58} studies in our review assessed documented occurrence in a specific module of the electronic medical record, suggesting that despite the potential of this process measure, it is still at its dawn and likely linked with the features of the local health-care information system.

The timing of a conversation is another key process indicator to be monitored. Evidence of benefit is accumulating on initiating conversations well before the last days of life.⁷⁰ However, end-of-life conversations are hindered by several obstacles and interventions targeting only professionals may not be enough. When interventions target multiple stakeholders, they work at multiple levels by reducing professionals' uncertainties about the right time to start conversations and simultaneously increasing patients' and family caregivers' question-asking about end-of-life care, ^{62,63} that was found to play an essential prompting role. ⁷¹

End-of-life conversations should allow adequate time to process and digest the information that are given.³² Being in a rush is a well-known obstacle to conversations and professionals are recommended to take the time to listen to patients and create the conditions to help the conversation run smoothly.⁷² In settings such as the nursing homes where stays are usually long and patients and family are familiar with the professionals, brief but more frequent encounters that break information into multiple chunks may work better to promote understanding and, finally, shared decision-making.

Our meta-analytic findings suggest no overall effect of the interventions on any process indicators of conversations despite several positive individual study outcomes. The high heterogeneity may be responsible for these null results. Unfortunately, there was not enough power for subgroup analyses to explore whether results differed for groups such as whom was targeted, employment of system level strategies, number of sessions, setting, family involvement, and trial design. However, visual inspection of the meta-analyses plots shows that subgroups of studies have the same clear and positive direction of the intervention effect. Particularly, in the plot of documented occurrence, two^{52,53} explanatory studies found the intervention to be effective while the effect was unclear or border-significant in pragmatic studies. 31,54-58 This suggests that the interventions have best chances to demonstrate a beneficial effect in ideal settings, while implementation in real care circumstance needs improvement. In the plot of patientreported occurrence, interventions targeting multiple stakeholders^{49,52} reported greater effect compared to interventions involving only professionals.^{50,51} This prompts more situational awareness that serves better understanding of illness and prognosis.

Regardless of limited effects on communication process indicators, the interventions may have taught professionals what issues to discuss⁷³ and possibly improved their sensitivity to addressing emotions. Such skills need time and practice to develop, therefore these interventions may not immediately result in improved patient-reported quality of conversations. Moreover, training was often a few hours, one-time ^{55–57,64} or over a few weeks. ^{50,60} Link of interventions to outcomes was not an objective of the study, but unchanged anxiety ^{52,53,63,65} and increased depression only in one study ⁵⁹ should reassure professionals that conversations about the end of life may be emotional but probably not harmful, and encourage them to engage in such conversations. ^{52,53,63}

The setting of interventions is a further recognized measure to assess the quality of care that needs to be considered. 17 Only five 53,58-60,63 studies involved outof-hospital settings (i.e., heart failure, primary care, palliative care or cancer clinics). However, the way health care is being delivered is changing; developed countries are adopting community-centered approaches and boosting out-of-hospital care. 74-76 Moreover, increasing care transitions in the late stages of illness may be avoided by primary care teams as better positioned to initiate and conduct regular end-of-life conversations to guide continuous care.⁷⁷ None of the studies were conducted in nursing homes, which is, among the out-of-hospital settings, at the frontline of caring for frail older people at the end of life and their family.⁷⁸ Only one⁵⁶ study involved patients with dementia and only in three 60,62,65 studies the intervention targeted family caregivers. When family caregivers are involved in end-of-life conversations, the likelihood to limit or withdraw life-sustaining treatments increases and it facilitates transitioning toward palliative-oriented care.^{8,79} Interventions to promote end-of-life conversations should be introduced and tested in particular in community settings and involve family caregivers.

Strengths and Limitations

Our findings should be read considering the lack of a common definition of end-of-life conversations, although this offers a picture of current practice. Second, documented occurrence of conversations may not reflect actual discussions or the provision of goal-concordant care. However, the outcomes were informed by several sources of information, including patients', family caregivers' or professionals' questionnaires or interviews, and videotaped or audio-recorded consultations in addition to medical records, in an effort to capture the nuances and content of end-of-life conversations beyond a simple binary outcome of occurrence. Third, we did not collect data about professionals' communication skills which are critical to establish trusting relationships that in turn may affect process and care outcome. Last, we did not have enough power to explore the subgroup effects of interventions aimed to promote end-of-life conversations in a specific setting or when family was involved.

Conclusions

This systematic review and meta-analysis found no effect of interventions aimed to promote end-of-life conversations on occurrence, quality, timing, frequency and duration of these conversations. Nevertheless, we found indications for interventions targeting multiple stakeholders to promote earlier and more comprehensive conversations; future study should identify effective elements. Overall, studies were mostly pragmatic even if settings involved, resources, flexibility in delivering the intervention and follow up modalities were mostly explanatory. This suggests implementation of interventions can be improved. Considerable heterogeneity renders conclusions tentative. Future research should examine the effect of interventions aimed to promote end-of-life conversations by employing standardized process indicators that represent the natural, basic starting point of evaluation to elicit implementation challenges or failure, and trigger improvement.

Author Contributions

BA, JTS, PB, PDG, SG, VD, and YA. For the purposes of authorship, Prof. Di Giulio and Prof. van der Steen contributed equally. Dr Gonella had full access to all the data in the study and takes responsibility for the integrity of the data and the accuracy of the data analysis. Conceptualization: Di Giulio, Dimonte, Gonella. Methodology: Di Giulio, Gonella, van der Steen. Software: Berchialla. Validation: Gonella. Formal analysis: Berchialla. Investigation: Arnone, Gonella. Resources: Arnone, Gonella. Data curation: Gonella. Writing—Original draft: Di Giulio, Gonella, van der Steen. Writing—Review & Editing: Albanesi, Arnone, Berchialla, Dimonte. Visualization: Gonella. Supervision: Di Giulio, van der Steen. Project administration: Gonella.

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This supplementary material is provided by the authors to give readers additional information about the system-

Appendix

atic review. Table A3 - Quality appraisal of the studies included in the systematic review according to the QualSyst critical Table A7 - Applicability of the interventions assessed in the studies included in the systematic review according to Figure A1 – Funnel plots of interventions aimed at promoting end-of-life conversations and the main outcomes of interests: documented occurrence in clinical records, patient-reported occurrence, and patient-reported Figure A2. Subgroup analyses of interventions aimed at promoting documented occurrence of end-of-life con-

Appendix 1

Tables A1-A7, Figures A1 and A2

Table A1 PRISMA Checklist

Section and Topic	Item#	Checklist Item	Location Where Item is Reported
Title			
Title	1	Identify the report as a systematic review.	Title page
Abstract		,	• 0
Abstract	2	See the PRISMA 2020 for Abstracts checklist.	Abstract, page 1
Introduction			
Rationale	3	Describe the rationale for the review in the context of existing knowledge.	Page 2
Objectives	4	Provide an explicit statement of the objective(s) or question(s) the review addresses.	Page 3
Methods			
Eligibility criteria	5	Specify the inclusion and exclusion criteria for the review and how studies were grouped for the syntheses.	Page 3
Information sources	6	Specify all databases, registers, websites, organizations, reference lists and other sources searched or consulted to identify studies. Specify the date when each source was last searched or consulted.	Page 3
Search strategy	7	Present the full search strategies for all databases, registers and websites, including any filters and limits used.	Page 3, Appendix 2
Selection process	8	Specify the methods used to decide whether a study met the inclusion criteria of the review, including how many reviewers screened each record and each report retrieved, whether they worked independently, and if applicable, details of automation tools used in the process.	Page 3

$Interventions\ to\ Promote\ End-of-Life\ Conversations\ in\ Clinical\ Encounters$

Table A1 Continued

Section and Topic	Item#	Checklist Item	Location Where Item is Reported
Data collection process	9	Specify the methods used to collect data from reports, including how many reviewers collected data from each report, whether they worked independently, any processes for obtaining or confirming data from study investigators, and if applicable, details of automation tools used in the	Page 3
Data items	10a	process. List and define all outcomes for which data were sought. Specify whether all results that were compatible with each outcome domain in each study were sought (e.g., for all measures, time points, analyses), and if not, the methods used to decide which results to collect.	Page 4
	10b	List and define all other variables for which data were sought (e.g., participant and intervention characteristics, funding sources). Describe any assumptions made about any missing or unclear information.	Page 3
Study risk of bias assessment	11	Specify the methods used to assess risk of bias in the included studies, including details of the tool(s) used, how many reviewers assessed each study and whether they worked independently, and if applicable, details of automation tools used in the process.	Page 3
Effect measures	12	Specify for each outcome the effect measure(s) (e.g., risk ratio, mean difference) used in the synthesis or presentation of results.	Page 4
Synthesis methods	13a	Describe the processes used to decide which studies were eligible for each synthesis (e.g., tabulating the study intervention characteristics and comparing against the planned groups for each synthesis [item #5]).	Page 4
	13b	Describe any methods required to prepare the data for presentation or synthesis, such as handling of missing summary statistics, or data conversions.	Page 4
	13c	Describe any methods used to tabulate or visually display results of individual studies and syntheses.	Page 4
	13d	Describe any methods used to synthesize results and provide a rationale for the choice(s). If meta—analysis was performed, describe the model(s), method(s) to identify the presence and extent of statistical heterogeneity, and software package(s) used.	Page 4
	13e	Describe any methods used to explore possible causes of heterogeneity among study results (e.g., subgroup analysis, meta-regression).	Page 4
	13f	Describe any sensitivity analyses conducted to assess robustness of the synthesized results.	Page 4
Reporting bias assessment	14	Describe any methods used to assess risk of bias due to missing results in a synthesis (arising from reporting biases).	Page 4

Table A1 Continued

Section and Topic	Item#	Checklist Item	Location Where Item is Reported
Certainty assessment	15	Describe any methods used to assess certainty (or confidence) in the body of evidence for an outcome.	Page 3
Results Study selection	16a	Describe the results of the search and selection process, from the number of records identified in the search to the number of studies included in the review, ideally using a flow diagram.	Page 4, Fig. 1
	16b	Cite studies that might appear to meet the inclusion criteria, but which were excluded, and explain why they were excluded.	Fig. 1, Table A2 (Appendix)
Study characteristics	17	Cite each included study and present its characteristics.	Page 4, Table 1
Risk of bias in studies	18	Present assessments of risk of bias for each included study.	Table 1, Table A3 (Appendix), page 4
Results of individual studies	19	For all outcomes, present, for each study: (a) summary statistics for each group (where appropriate) and (b) an effect estimate and its precision (e.g., confidence/credible interval), ideally using structured tables or plots.	Page 20-21, Table 1
Results of syntheses	20a	For each synthesis, briefly summarize the characteristics and risk of bias among contributing studies.	Page 20-21, Fig. 2, Fig. A1 (Appendix)
	20b	Present results of all statistical syntheses conducted. If meta-analysis was done, present for each the summary estimate and its precision (e.g., confidence/credible interval) and measures of statistical heterogeneity. If comparing groups, describe the direction of the effect.	Page 20-21, Fig. 2
	20с	Present results of all investigations of possible causes of heterogeneity among study results.	Page 20, Fig. A2 (Appendix)
	20d	Present results of all sensitivity analyses conducted to assess the robustness of the synthesized results.	Page 20
Reporting biases	21	Present assessments of risk of bias due to missing results (arising from reporting biases) for each synthesis assessed.	Table 1, Fig. A1 (Appendix)
Certainty of evidence	22	Present assessments of certainty (or confidence) in the body of evidence for each outcome assessed.	Page 21, Table 1, Fig. 2, Table A7 (Appendix)
Discussion Discussion	23a	Provide a general interpretation of the results in the context of other evidence.	Page 21
	23b	Discuss any limitations of the evidence included in the review.	Page 22
	23с	Discuss any limitations of the review processes used.	Page 22
	23d	Discuss implications of the results for practice, policy, and future research.	Page 22
Other Information Registration and protocol	24a	Provide registration information for the review, including register name and registration number, or state that the review was not registered.	Page 3
	24b	Indicate where the review protocol can be accessed, or state that a protocol was not prepared.	Page 3
	24c	Describe and explain any amendments to information provided at registration or in the protocol.	Page 3

Table A1 Continued

Section and Topic	Item#	Checklist Item	Location Where Item is Reported
Support	25	Describe sources of financial or non- financial support for the review, and the role of the funders or sponsors in the review.	Page 23
Competing interests	26	Declare any competing interests of review authors.	Page 23
Availability of data, code and other materials	27	Report which of the following are publicly available and where they can be found: template data collection forms; data extracted from included studies; data used for all analyses; analytic code; any other materials used in the review.	Available to authors

From: Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. BMJ 2021;372:n71. doi: 10.1136/bmj.n71.

 $Table\,A2$ Excluded Articles Based on Full Text Review

Author(s), year	Reason(s) for Exclusion	Reference
Alexander et al., 2006	No data on process indicators of end-of-life conversations	Alexander SC,Keitz SA, Sloane R, et al. A controlled trial of a short course to improve residents' communication with patients at the end of life. <i>Acad Med.</i> 2006; 81(11):1008-12. doi: 10.1097/01. ACM.0000242580.83851.ad.
Bernacki et al., 2019	No data on process indicators of end-of-life conversations	Bernacki R, Paladino J, Neville BA, et al. Effect of the Serious Illness Care Program in outpatient oncology: A cluster randomized clinical trial. <i>JAMA Intern Med.</i> 2019; 179(6):751-759. doi: 10.1001/ jamainternmed.2019.0077.
Chikhladze et al., 2016	No intervention study	Chikhladze N, Janberidze E, Velijanashvili M, et al. Mismatch between physicians and family members views on communications about patients with chronic incurable diseases receiving care in critical and intensive care settings in Georgia: a quantitative observational survey. <i>BMC Palliat Care</i> . 2016; 22;15:63. doi: 10.1186/s12904-016-0135-2.
Curtis et al., 2012	No data on process indicators of end-of-life conversations	Curtis JR,Ciechanowski PS, Downey L, et al. Development and evaluation of an interprofessional communication intervention to improve family outcomes in the ICU. <i>Contemp Clin Trials</i> . 2012; 33 (6):1245-54. doi: 10.1016/j.cct.2012.06.010.
Fakhiri et al., 2016	No intervention study	Fakhri S,Engelberg RA, Downey L, et al. Factors affecting patients' preferences for and actual discussions about End-of-Life care. <i>J Pain Symptom Manage</i> . 2016; 52(3):386-94. doi: 10.1016/j. jpainsymman.2016.03.012.
Gradwohl et al., 2020	Intervention part of a multi-faceted programme with the effect of the individual component of the intervention related to end-of-life communication not clearly recognizable and assessable	Gradwohl K, Wood GJ, Clepp RK, et al. Preventing Readmissions through Effective Partnerships-Communication and Palliative Care (PREP-CPC): a multisite intervention for encouraging goals of care conversations for hospitalized patients facing serious Illness. <i>Am J Hosp Palliat Care</i> . 2020, 37(8):582-588. doi: 10.1177/1049909119891996.
Izumi et al., 2019	No data on process indicators of end-of-life conversations	Izumi S, Burt M, Smith J, et al. Enhancing Advance Care Planning conversations by nurses in a bone marrow transplantation unit. <i>Oncol Nurs Forum.</i> 2019; 46 (3):288-297. doi: 10.1188/19.ONF.288-297.
Maries et al., 2018	No data on process indicators of end-of-life conversations	Maries MM, Kryworuchko J, Vininder Kour B, et al. A randomized controlled trial of a serious illness communication workshop for nurses in critical care. <i>Can J Crit Care Nurs.</i> 2018; 29(2): 58-58.
Modes et al., 2019	Secondary analysis which merged the intervention and the control groups	Modes ME, Engelberg RA, Downey L, et al. Did a Goals- of-Care discussion happen? Differences in the occurrence of Goals-of-Care discussions as reported by patients, clinicians, and in the electronic health

Table A2 Continued

Author(s), year	Reason(s) for Exclusion	Reference
Nedjat-Haiem et al., 2019	No data on process indicators of end-of-life conversations	record. JPain Symptom Manage. 2019; 57(2):251-259. doi: 10.1016/j.jpainsymman.2018.10.507. Nedjat-Haiem FR,Cadet TJ, Amatya A, et al. Efficacy of motivational interviewing to enhance advance directive completion in latinos with chronic illness: a randomized controlled trial. Am J Hosp Palliat Care. 2019; 36(11):980-992. doi: 10.1177/
Pajka et al., 2021	No data on process indicators of end-of-life conversations	1049909119851470. Pajka SE, Hasdianda MA, George N, et al. Feasibility of a brief intervention to facilitate advance care planning conversations for patients with life-limiting Illness in the Emergency Department. <i>J Palliat Med.</i> 2021; 24 (1):31-39. doi: 10.1089/jpm.2020.0067.
Paladino et al., 2020	No data on process indicators of end-of-life conversations	Paladino J, Koritsanszky L, Neal BJ et al. Effect of the Serious Illness Care Program on health care utilization at the end of life for patients with cancer. <i>J Palliat Med.</i> 2020; 23(10):1365-1369. doi: 10.1089/jpm.2019.0437.
Paladino et al., 2020	No data on process indicators of end-of-life conversations	Paladino J, Kilpatrick L, O'Connor N, et al. Training clinicians in serious illness communication using a structured guide: evaluation of a training program in three health systems. <i>J Palliat Med.</i> 2020; 23(3):337-345. doi: 10.1089/jpm.2019.0334.
Sanchez et al., 2018	Commentary	Sanchez R, Mateo KF. Survey-based priming intervention linked to improved communication with the seriously Ill. <i>J Clin Outcomes Manage</i> . 2018; 25 (7):300-303.
Song et al., 2010	Unclear clinical condition of advanced chronic or terminal illness	Song MK, Donovan HS, Piraino BM, et al. Effects of an intervention to improve communication about end-of-life care among African Americans with chronic kidney disease. <i>Appl Nurs Res.</i> 2010; 23(2):65-72. doi: 10.1016/j.apnr.2008.05.002.
Steinhauser et al., 2008	No data on process indicators of end-of-life conversations	Steinhauser KE, Alexander SC, Byock IR et al. Do preparation and life completion discussions improve functioning and quality of life in seriously ill patients? Pilot randomized control trial. <i>J Palliat Med.</i> 2008; 11 (9):1234-40. doi: 10.1089/jpm.2008.0078.
Wentlandt et al., 2012	No data on process indicators of end-of-life conversations	Wentlandt K, Burman D, Swami N, et al. Preparation for the end of life in patients with advanced cancer and association with communication with professional caregivers. <i>Psychooncology</i> . 2012; 21(8):868-76. doi: 10.1002/pon.1995.

Table A3 Quality Appraisal of the Studies Included in the Systematic Review According to the QualSyst Critical Appraisal Criteria^a

		~	, 11							<u>∘</u> ~	•	11				
Author(s), year	CI. Question / Objective Suffi Ciently Described?	C2. Study Design Evident and Appropriate?	C3. Method of Subject / Comparison Group Selection or Source of Information/ Input Variables Described and Appropriate?	and Comparison Group characteristics Sufficiently	C5. Was Random Allocation Described?	C6. Was Blinding of Investigators Reported?	C7. Was Blinding of Subjects Reported?	C8. Outcome Measures Well Defined and Robust?		C10. Analytic Methods Described and Appropriate?	C11. Some Estimate of Variance Reported for the Main Results?	C12. Controlled for Confounding?		C14. Conclusions Supported by the Results?	Quality Rating (Total Sum/ Total Possible Sum)	Quality Rating
Au et al., 2012	2	2	2	2	1	2	N/A	2	2	2	2	2	2	2	0.96	Н
Bickell et al., 2020	2	1	2	2	1	2	N/A	2	2	2	2	2	2	2	0.92	Н
Clayton et al., 2007		2	2	1	2	2	N/A	2	2	2	2	2	2	2	0.96	Н
Connors et al., 1995	2	2	2	2	1	2	N/A	2	2	2	2	2	2	2	0.96	Н
Curtis et al., 2013		2	2	2	1	2	N/A	2	2	2	2	2	2	2	0.96	Н
Curtis et al., 2018		2	2	1	2	2	N/A	2	2	2	2	2	2	2	0.96	Н
Doorenbos et al., 2016 Epstein et al.		2	2	2	1	2	N/A N/A	2	2	2	2	2	2	2	0.88	M H
, 2017 Hanson	2	1	2	1	N/A	N/A	N/A	2	1	2	0	0	2	2	0.68	L
et al., 2017 Henselmans		2	2	2	1 1	2	N/A	2	2	2	2	2	2	2	0.96	Н
et al., 2020 Lakin et al.,		1	1	2	0	2	N/A	2	1	2	0	0	2	2	0.65	L
2017 Lautrette	2	2	2	2	2	2	N/A	2	2	2	2	0	2	2	0.92	Н
et al., 2007 Nakajima	2	1	1	2	N/A	N/A	N/A	1	0	2	2	0	2	2	0.68	L
et al., 2015 Paladino et al., 2019	2	2	2	2	1	2	N/A	2	2	2	2	2	2	2	0.96	Н
et al., 2019 Pollak et al. 2019		1	2	2	2	0	N/A	2	1	2	2	0	2	2	0.77	M
Reinke et al., 2017	2	2	2	2	N/A	0	N/A	2	1	2	2	2	2	2	0.88	M
Shorr et al., 2000	2	2	2	2	N/A	N/A	N/A	2	1	2	2	0	2	2	0.86	M
Walczak et al., 2017	2	2	2	2	2	0	N/A	2	1	2	2	2	2	2	0.88	M

aKmet LM, Cook LS, Lee RC. Standard quality assessment criteria for evaluating primary research papers from a variety of fields. Alberta Heritage Foundation for Medical Research; 2004. doi: 10.7939/R37M04F16Abbreviations. C, criteria; H, high; L, low; M, moderate. Note: This Qualsyst tool comprises of 14 items which were scored on the degree to which the specific criteria were met ("fully" = 2, "partially" = 1, "not at all" = 0). Items not relevant for a particular study design were marked "N/A" and excluded from the calculation of the summary score. A summary score was calculated for each paper by summing the score obtained across relevant items and dividing by the total possible score after removing any criteria that were not relevant. According to the summary score, studies were classified as high (>0.9), moderate (0.7-0.9) or low quality (<0.7).

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Table A4 **Data Collection Tools and Timing in the Included Studies**

Data concessor 1000 and 111115 in the metades studies						
Data Collection Tools	Data Collection Timing					
Medical records $(n = 7)^{31,52-58}$ Patient questionnaires $(n = 6)^{49,50,52,53,59,61}$ Audio-recorded consultations 60,62,63 Videotaped consultations 64 Family questionnaires 59 Professionals questionnaires 59 Patients interviews 51 Family caregivers interviews 51 Clocked consultations 65	A few days after the intervention ⁵⁰ One or two weeks after the intervention ^{49,51-53,61} Six to ten months after the intervention ^{50,59}					

Table A5 Interventions Aimed at Promoting End-of-Life Conversations and Their Target

interventions Aimed at Fromoung End-of-Life Conversations and Their Target									
Interventions Targeting Both Patients and Professionals	Interventions Targeting Only Professionals	Interventions Targeting Only Patients	Interventions Targeting the System						
Coaching $(n = 7)^{50,53,55,56,58,60,62}$ Written resources such as question prompts list, communication guide, and information leaflet $(n = 6)^{55,60,62-65}$ Patient-specific information form about end-of-life care preferences $(n = 4)^{31,49,51,53}$ Videos with educational or supportive purposes $(n = 3)^{60,62,54}$ Patient-specific communication tips form 52,53	Lectures $(n = 6)^{50,54,55,57,59,64}$ Role play $(n = 6)^{50,57-60,64}$ Individual- or small group- based reflective discussions $(n = 5)^{54,57,59,60,64}$ Cues to use communication aids $(n = 3)^{62,63,65}$ Provision of a prognostic estimate ⁵¹ Palliative care consultation ⁵⁴	Disclosure of more detailed information about clinical conditions ⁶¹ Face-to-face meeting ⁶²	Email remainders, alerts, or administrative prompts ^{56,57} Electronic documentation template for conversations ^{55,58} Trained nurse who facilitated conversations ^{51,58}						

$Table\,A6$ Patients- and Family Caregivers-Related Care Outcomes Reported in the Included Studies

Patients	Family Caregivers
Hospitalization or readmission $(n = 4)^{50,54,56,a,60}$	Depression ⁵⁹
Depression $(n = 4)^{52,53,59,0,05,0}$	Satisfaction with information received 65
Anxiety $(n = 4)^{52,53,63,65,c}$	
Aggressive treatments $(n = 3)^{50,51,60}$	
Palliative care or hospice referral $(n = 3)^{53,56,58}$	
Quality of life or well-being $(n = 3)^{59,62,64}$	
Length of in-hospital or intensive care units (ICU) stay $(n = 2)^{51,56}$	
Emergency department visits $(n = 2)^{50,60}$	
ICU admission $(n = 2)^{50,54}$	
In-hospice stay (n = 2) 50,58	
Palliative care consultations 54,d	
Post-traumatic stress disorder 65,c	
Symptom assessment 54,d	
Time in pain before death ⁵¹	
Assessment of spiritual beliefs ⁵⁴	
Satisfaction and uncertainty about medical decisions ⁶⁴	
Satisfaction with communication ⁶³	
Achievement of information needs ⁶³	
Completion of advance directives \$1,51,53,57	
Documentation of advance care planning ⁵⁶	

Note. Four 54,56,59,65 studies found significant changes with mixed results. Three 54,56,59 targeted only professionals, one 65 multiple stakeholders. a Decrease in overall (P < 0.001) and 30-day readmission rates (P = 0.009) with improvement of the content without changes in the occurrence of conversations. 56 bIncreased patients' depression (P = 0.006) without improvement in patient, family caregivers, or clinician-reported reported quality of conversations. 59 Correcased patients' post-traumatic stress disorder (P = 0.01), anxiety (P = 0.02), and depression (P = 0.003) with longer conversations 65 dIncreased rate of palliative care consultation (P = 0.026) but reduced screening of constipation (P = 0.041) in presence of increased occurrence of conversations in clinical records. 54

 ${\it Table\,A7}$ Applicability of the Interventions Assessed in the Studies Included in the Systematic Review According to the PRECIS-2 Tool $^{\rm a}$

Author(s), year	D1. Eligibility Criteria - Who Is Selected to Participate in the Trial?	D2. Recruitment - How are Participants Recruited into the Trial?	D3. Setting - Where is the Trial Being Done?	D4. Organisation - What Expertise and Resources are Needed to Deliver the Intervention?	D5. Flexibility (delivery) - How Should the Intervention be delivered?	D6. Flexibility (Adherence) - What Measures are in Place to Ensure Participants Adhere to the Intervention?	D7. Follow up - How Closely are Participants Followed up?	D8. Primary Outcome - How Relevant is it to Participants?	D9. Primary Analysis - To What Extent are all Data Included?	Mean Applicability Rating (Total Sum/9)	Applicability
Au et al., 2012	4	4	4	4	4	5	3	5	5	4.2	Mostly pragmatic
Bickell et al., 2020	4	5	5	3	5	5	4	5	2	4.2	Mostly pragmatic
Clayton et al., 2007	4	5	5	4	4	5	2	5	5	4.4	Mostly pragmatic
Connors et al., 1995	2	5	5	3	5	5	3	5	3	4	Mostly pragmatic
Curtis et al., 2013	4	4	5	3	2	5	4	5	2	3.7	Mostly explanatory
Curtis et al., 2018	5	3	5	3	2	5	4	5	2	3.7	Mostly explanatory
Doorenbos et al., 2016	2	3	2	3	2	5	4	4	5	3.3	Mostly explanatory
Epstein et al., 2017	4	2	5	3	3	5	2	5	5	3.7	Mostly explanatory
Hanson et al., 2017	5	5	2	2	4	3	5	5	5	4.0	Mostly pragmatic
Henselmans et al., 2020	5	5	5	4	2	5	4	5	5	4.4	Mostly pragmatic
Lakin et al., 2017	5	2	5	4	2	5	4	5	5	4.1	Mostly pragmatic
Lautrette et al., 2007	5	5	5	3	2	5	3	5	4	4.1	Mostly pragmatic
Nakajima et al., 2015	3	3	2	5	5	5	4	5	5	4.1	Mostly pragmatic
Paladino et al., 2019	5	2	5	4	3	5	5	5	5	4.3	Mostly pragmatic
Pollak et al. 2019	5	5	1	4	5	5	5	5	4	4.3	Mostly pragmatic
Reinke et al., 2017	4	4	4	4	4	5	5	5	5	4.4	Mostly pragmatic
Shorr et al., 2000	5	5	1	4	5	5	5	5	5	4.4	Mostly pragmatic
Walczak et al., 2017	5	5	5	4	2	5	4	5	5	4.4	Mostly pragmatic
All studies	4.2	4.0	3.9	3.6	3.4	4.9	3.9	4.9	4.3	4.0	Four domains <4 , five domains ≥ 4

Abbreviations: D, domain; PRECIS-2, PRagmatic Explanatory Continuum Indicator Summary version 2.

Note: This PRECIS-2 tool is focused on the issue of applicability of study results and comprises of nine domains. Each domain was scored on a 5-point scale (1 = very explanatory to 5 = very pragmatic). When with insufficient information, a score of 3 was assigned. A mean score across studies for each domain and a mean applicability rating for each study was calculated. Domains and studies were classified as mostly explanatory (<4) or mostly pragmatic (≥ 4) according to the mean domain score and the mean applicability rating, respectively. Values with decimals were rounded to the lower integer.

^aLoudon K, Treweek S, Sullivan F, Donnan P, Thorpe KE, Zwarenstein M. The PRECIS-2 tool: designing trials that are fit for purpose. J Clin Epidemiol. 2017;88:113-121. doi:10.1016/j.jclinepi.2017.06.001.

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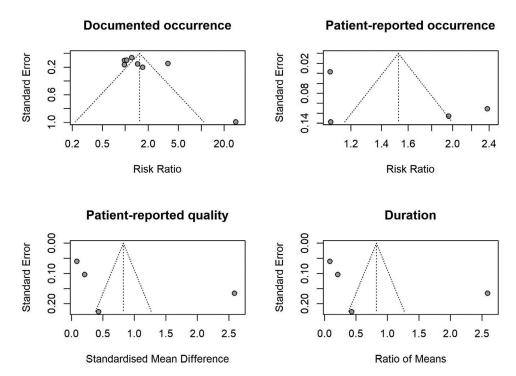


Fig. A1. Funnel plots of interventions aimed at promoting end-of-life conversations and the main outcomes of interests: documented occurrence in clinical records, patient-reported occurrence, patient-reported quality, and duration of end-of-life conversations.

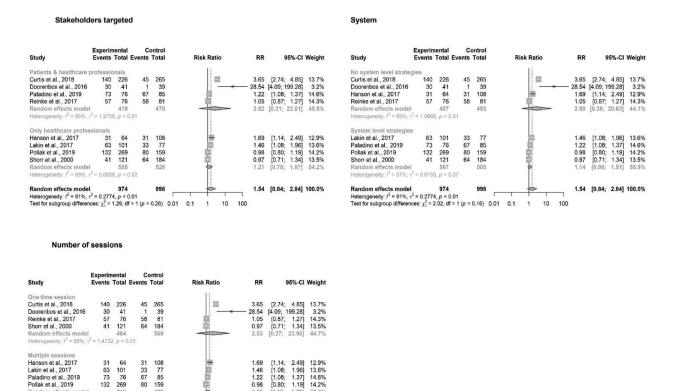


Fig. A2. Subgroup analyses of interventions aimed at promoting documented occurrence of end-of-life conversations in clinical records: stakeholders targeted, system strategies, and number of sessions. CI, Confidence interval; df, degrees of freedom; RR, Risk ratio.

1.54 [0.84; 2.84] 100.0%

1 10 100

Appendix 2. Search strategies and hits September 30, 2021

1. Pubmed.

Search	Query	Items
#5	(#1 AND #2 AND #3) Filters: Clinical Trial, Controlled Clinical Trial, Observational	483
	Study, Pragmatic Clinical Trial, Randomized Controlled Trial	
#4	#1 AND #2 AND #3	10,989
#3	("Documentation" [Mesh] OR "Quality Improvement" [Mesh] OR occurrence* [Title/	4,695,319
	Abstract] OR timing [Title/Abstract] OR duration [Title/Abstract] OR quality	
	[Title/Abstract] OR "accessibility document*" [Title/Abstract] OR frequen*[Title/	
	Abstract])	
#2	("Communication" [Mesh] OR "Health Communication" [Mesh] OR "Advance Care	5,617,810
	Planning" [Mesh] OR "Decision Making" [Mesh] OR information [Title/Abstract] OR	
	conversation* [Title/Abstract] OR talk*[Title/Abstract] OR discuss*[Title/Abstract]	
	OR new*[Title/Abstract] OR dialogue*[Title/Abstract])	
#1	("Death" [Mesh] OR "Terminal Care" [Mesh] OR "Terminally Ill" [Mesh] OR "end-of-	221,676
	life"[Title/Abstract] OR "end of life" [Title/Abstract])	

2. EBSCO CINAHL.

Search ID	Search Terms	Search Options	Actions
S5	S4	Limiters - Peer Reviewed; Exclude MEDLINE records; Publication Type: Clinical Trial, Randomized Controlled Trial	178
		Search modes - Boolean/Phrase	
S4	S1 AND S2 AND S3	Search modes - Boolean/Phrase	23,166
S3	MH ("documentation+" OR "clinical documentation improvement" OR "quality improvement+" OR "access to information+") OR AB (documentation OR occurrence* OR frequen* OR timing OR duration OR quality OR "accessibility document") OR TI (documentation OR occurrence* OR frequen* OR timing OR duration OR quality OR "accessibility document")	Search modes - Boolean/Phrase	1,099,385
S2	MH ("communication+" OR "advance care planning" OR "decision making, patient+" OR "decision making, family") OR AB (communication OR "health communication" OR "advance care planning" OR information OR conversation* OR talk* OR discuss* OR new* OR dialogue* OR "decision making") OR TI (communication OR "health communication" OR "advance care planning" OR information OR conversation* OR talk* OR discuss* OR new* OR dialogue* OR "decision making")	Search modes - Boolean/Phrase	1,775,546
S1	MH "terminal care+" OR AB (death OR dying OR "terminally ill" OR "end of life" OR "end-of-life") OR TI (death OR dying OR "terminally ill" OR "end of life" OR "end-of-life")	Search modes - Boolean/Phrase	270,753

3. EBSCO PsycINFO.

Search ID	Search Terms	Search Options	Actions
S5	S4	Limiters - Methodology: CLINICAL TRIAL, EMPIRICAL STUDY, QUANTITATIVE STUDY; Exclude Dissertations Search modes - Boolean/Phrase	2,004
S4	S1 AND S2 AND S3	Search modes - Boolean/Phrase	8,025
S3	AB (documentation OR "quality improvement" OR occurrence* OR frequen* OR timing OR duration OR quality OR "accessibility document") OR TI (documentation OR "quality improvement" OR occurrence* OR frequen* OR timing OR duration OR quality OR "accessibility document")	Search modes - Boolean/Phrase	734,569
S2	DE ("Interpersonal Communication" OR "Verbal Communication" OR "communication" OR "Treatment Planning" OR "Decision Making") OR AB ("advance care planning" OR information OR conversation* OR talk* OR discuss* OR new* OR dialogue* OR "decision making") OR	Search modes - Boolean/Phrase	1,845,313

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Continued

Search ID	Search Terms	Search Options	Actions
S1	TI ("advance care planning" OR information OR conversation* OR talk* OR discuss* OR new* OR dialogue* OR "decision making") DE ("palliative care" OR "terminally ill patients") OR AB (death OR "terminal care" OR "terminally ill" OR "end of life" OR "end-of-life") OR TI (death OR "terminal care" OR "terminally ill" OR "end of life" OR "end-of-life")	Search modes - Boolean/Phrase	109,507
4. SCOPUS.			
Query			Items
TITLE-ABS-KEY (death OR terminal OR end) AND (communication* OR discussion* OR conversation*) AND (documentation OR quality OR frequen*)			