

# The context, components and mechanisms of action for early and/or integrated palliative care in adult oncology: a systematic review with assessment of transferability into resource-limited settings

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## Abstract

**Background:** Clinical trials have demonstrated that palliative care (PC) provided early and alongside cancer treatment can alleviate symptoms and enhance the quality of life. This review aimed to synthesise the existing evidence for an early integrated PC in adult oncology with a specific focus on context, components, mechanisms of action and transferability into low-income settings.

**Methods:** Systematic review of six databases 1967–2025, identifying randomised controlled trials evaluating early and/or integrated PC in adult oncology. Quality appraisal used the Mixed Methods Appraisal Tool (MMAT). Data were synthesised via narrative and intervention syntheses. Context, components, mechanisms of action and outcomes of PC were mapped in a logic model. Transferability was assessed by the TRANSFER approach.

**Results:**  $N = 46$  papers (2010–25) were retained, with  $n = 40$  (93%) meeting the quality assessment criteria, fulfilling 4/5 MMAT criteria, with only  $n = 6$  (13%) from the global south. Nurses were the most common PC providers, delivering patient communication and symptom management, either within multidisciplinary teams, in doctor–nurse dyads or as sole providers of PC. The term ‘early’ denoted initiation of PC: (a) within 8 weeks of an advanced cancer diagnosis, (b) linked to a cancer treatment phase or (c) based on an estimated prognosis of 6–24 months. Three models of integration were identified: (a) co-located concurrent care (basic integration of services), (b) interdisciplinary collaboration (e.g. team meetings and shared decision making), (c) fully embedded/unified care (e.g. co-rounding models or primary PC by oncology providers). With respect to transferability to low-resource settings, while the core principles are relevant, direct replication is limited by the evidence being from high-resource contexts. Models emphasising structured protocols, task shifting to nurses and simplified, adaptable interventions are considered more feasible, whereas complex, specialist team-dependent models raise moderate concerns for transferability.

**Conclusion:** Locally appropriate models are urgently needed to meet the needs of patients and families in the global south, which faces the greatest cancer burden.

**Keywords:** *early, integrated, palliative care, oncology, cancer, models, resource-limited settings*

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## Introduction

Cancer was the second leading cause of death globally in 2020 and is expected to become the first by 2060 [1]. Cancer places a heavy burden on patients, families and healthcare systems, including symptoms such as pain, depression, the financial strain of treatment coupled with lost family income and caregiving demands and unplanned admissions [2, 3]. Economic and lifestyle changes, such as higher tobacco use, poor nutrition and physical inactivity, are driving cancer incidence increases in countries undergoing economic transition [4]. By 2060, four-fifths of global deaths with serious health-related suffering will occur in low- and middle-income countries (LMICs), where access to palliative care (PC) is poorest [1].

Despite evidence that PC improves outcomes and saves costs [5–8], currently specialist PC services only reach an estimated 14% of those who would benefit [9]. PC is an essential health service under Universal Health Coverage. Traditionally, PC has been closely linked to end-of-life care [10]. However, evidence suggests significant benefits of introducing PC early and/or alongside disease-oriented treatment. Systematic reviews of early or integrated PC report improvements in quality of life, symptom burden (including pain, depression and anxiety) and patient satisfaction with care [5, 6]. Furthermore, such integration is associated with reduced non-beneficial end-of-life treatments, lower healthcare costs and, in some cases, prolonged patient survival [7, 8].

The American Society of Clinical Oncology has defined early PC as starting within 8 weeks of diagnosis [11]. It is also defined by the clinical milestones in the disease course – for instance, at the time metastatic disease is identified [12] or when first-line palliative chemotherapy begins [13]. This variability reflects the evolving understanding that ‘early’ integration is not only about time but also about the point in the illness trajectory where PC can proactively address unmet needs alongside potentially disease-modifying treatment.

Integration of health care is defined as any ‘managerial or operational changes to health systems to bring together inputs, delivery, management and organisation of particular service functions as a means of improving coverage, access, quality, acceptability and cost-effectiveness’ [14]. Integrated PC is defined as ‘coordinated and continuous care across the different health organisations, levels of care and PC providers’ [15]. Within oncology care, integration has been defined as the systematic incorporation of PC principles and services into the overall care of cancer patients [16].

Existing reviews have effectively identified core components of integrated models, implementation barriers and economic impacts in various contexts [5, 17, 18]. Others have focused on specific themes such as the role of multidisciplinary teams [19] and the necessity of provider training [20]. Notably, some work has begun to explore access challenges in particular settings, including rural environments where resource constraints are often most acute [21]. Despite these contributions, significant gaps remain in understanding how PC is integrated into oncology, the mechanisms by which it provides its benefits, and how these findings can be translated into resource-limited settings, where the greatest need is. While previous reviews establish that integration works, a synthesis of how it works (i.e. through specific components, contextual factors and causal mechanisms) and transferability to low-resource settings is lacking.

This review aimed to identify, critically appraise and synthesise the existing evidence for integrated and/or early PC in adult oncology. The objectives were as follows: (a) to identify randomised controlled trials of early and/or integrated PC for oncology patients; (b) to appraise the study quality; (c) to map models focusing on context, intervention components and mechanisms of action and (d) to assess the transferability of evidence with particular attention to resource requirements.

## Methods

### Design

This systematic review utilised the PRISMA guidelines (Preferred Reporting Items for Systematic Reviews and Meta-Analysis) and the Guidance on the Conduct of Narrative Synthesis in Systematic Reviews [22, 23].

## Protocol registration

The protocol was registered on PROSPERO (CRD42023399630).

## Search strategy

### Databases

Six databases were searched: Embase, Scopus, CINAHL, MEDLINE, Global Health and the Cochrane Library. Manual search of relevant medical and nursing journals was undertaken. Our search range encompassed the period from 1967 (the inception of the modern PC movement with the establishment of the first modern hospice [24]) to January 2025.

### Search terms

The search was informed by prior reviews and included a combination of Medical Subject Headings and free-text terms. The search terms were structured into four main categories.

Firstly, terms related to PC: 'palliative medicine', 'palliative therapy', 'PC', 'terminal care', 'hospice care', 'palliat\*' and 'end of life care'.

Secondly, terms synonymous with 'cancer': 'cancer', 'oncolog\*', 'malignancy' and 'neoplasm'.

Thirdly, 'integration', 'early integration', 'integrated', 'early integrated', 'early systems integration', 'early integrat\*' and 'early'.

Finally, study design: the terms 'randomised controlled trials' and 'RCT' were included.

Within each category, individual terms were combined using the Boolean operator 'OR'. The four resulting sets were then combined using the Boolean operator 'AND' to retrieve articles relevant to all concepts.

Table 1 shows the detailed search terms for the MEDLINE database. The same strategy, with necessary adaptations, was applied to the other five databases.

### Eligibility criteria

Study inclusion criteria are reported in Table 2.

### Study selection and data extraction

Search results were uploaded to Covidence (<http://www.covidence.org>) and deduplicated before screening.

Firstly, title and abstract screening was conducted by the first reviewer (Nahla Gafer). Unclear cases were flagged for further review. Articles categorised as 'maybe' were discussed with the second reviewer (Nuhamin Gebre) to reach consensus.

Secondly, the first and second reviewers independently assessed full texts (approximately 60% of papers Nahla Gafer and 40% Nuhamin Gebre). Inter-rater agreement for full-text screening was 92%, with disagreements resolved through discussion with senior reviewers (Richard Harding and Matthew Maddocks).

A data extraction sheet was developed based on the Template for Intervention Description and Replication [25]. This 12-item reporting framework helps to ensure that interventions are described adequately for assessment or replication. Data extraction was performed by two independent investigators (Nuhamin Gebre and Khulood Alyami), followed by verification and completion by the first reviewer (Nahla Gafer) using Covidence and Google Sheets. Any discrepancies were resolved by consensus or escalated to Matthew Maddocks and Richard Harding.

Table 1. MEDLINE search strategy.

|     |                                                                                                                                                                                                                                                                                                                         |
|-----|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| 1.  | exp PC/                                                                                                                                                                                                                                                                                                                 |
| 2.  | palliative care.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]     |
| 3.  | exp Terminal Care/                                                                                                                                                                                                                                                                                                      |
| 4.  | terminal care.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]       |
| 5.  | palliative therapy.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]  |
| 6.  | palliative medicine.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms] |
| 7.  | supportive care.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]     |
| 8.  | end of life care.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]    |
| 9.  | hospice.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]             |
| 10. | 1 or 2 or 3 or 4 or 5 or 6 or 7 or 8 or 9                                                                                                                                                                                                                                                                               |
| 11. | exp Oncology Nursing/ or oncology.mp. or exp Medical Oncology/ or exp Radiation Oncology/                                                                                                                                                                                                                               |
| 12. | oncology.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]            |
| 13. | exp Neoplasms/                                                                                                                                                                                                                                                                                                          |
| 14. | neoplasm.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]            |
| 15. | cancer.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]              |
| 16. | 11 or 12 or 13 or 14 or 15                                                                                                                                                                                                                                                                                              |
| 17. | early.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]               |
| 18. | integrated.mp. [mp=title, book title, abstract, original title, name of substance word, subject heading word, floating sub-heading word, keyword heading word, organism supplementary concept word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier, synonyms]          |
| 19. | 17 or 18                                                                                                                                                                                                                                                                                                                |
| 20. | 10 and 16 and 19                                                                                                                                                                                                                                                                                                        |

**Table 2. Eligibility criteria according to population, intervention, control, outcome and study (PICOS) [28].**

| PICOS Element | Inclusion Criteria                                                                                                                                                               | Exclusion Criteria                                                                                                                                                                                                       |
|---------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Population    | Adult patients, 18 years or above cancer patients regardless the type and stage                                                                                                  | Aged under 18 years<br>>50% study population non-cancer and not reported separately                                                                                                                                      |
| Intervention  | PC, here it denotes holistic PC services delivered by at least two types of cadres trained/specialised in PC. It must either be described as early or integrated by the authors. | studies reporting solely palliative chemotherapy, palliative surgical interventions, or palliative radiotherapy<br>pure supportive care or best supportive care (especially when dealing with drug trials) was excluded* |
| Control       | Patients receiving concurrent oncology care, with or without a PC service.                                                                                                       |                                                                                                                                                                                                                          |
| Outcome       | Any patient-centred outcomes                                                                                                                                                     | Studies not reporting any patient-centred outcomes, e.g. health professional's outcomes                                                                                                                                  |
| Study         | Articles published from 1967 to Mar 2024<br>randomised controlled trials only**                                                                                                  | Systematic reviews, protocols, conference abstracts and proceedings                                                                                                                                                      |

\* Although supportive care has been defined as (the prevention and management of the adverse effects caused by treatment of life threatening and life limiting illnesses, including both physical and psychosocial symptoms, existential concerns and side effects across the entire continuum as well as the enhancement of rehabilitation and survivorship), [29] supportive care or best supportive care in clinical trials, are usually ill-defined and lack some components of care. In a systematic review less than a quarter of all studies had symptom management [30]. Likewise, patients too weak for cancer treatment are often given 'best supportive care', but this label doesn't always translate to helpful action. Such trials rarely report that patients had access to PC specialists or other support services, including nursing, social, financial and spiritual counselling [31]

\*\* Randomised controlled trials in this study denote any study that the author described as 'randomised' anywhere in the manuscript, except where it is misused to mean just a random sample. All types of randomised controlled trials (e.g. double-/single-blind or not blinded RCT, in addition to cluster RCT). Randomized controlled trials (RCTs) were chosen for their superior ability to reduce bias and generate high-quality evidence. The inclusion of non-RCTs, for example due to shortage of relevant literature was not applicable in this study, aligning with the guidelines outlined in the Cochrane Handbook [32]

Extracted variables included: study characteristics (author, year of publication, country, setting, country level of PC development using the WHO Global Atlas of PC [9] and aim of the study), population characteristics (type of cancer and disease stage), intervention details (type of PC service, description of how the intervention was delivered, team members who provided the intervention and their level of training or expertise, mode of delivery, timing for referral/intervention initiation, components and frequency of the intervention, materials used and barriers and facilitators); study design (type of RCT, recruitment period, assessment timeframes and method of data collection) and outcomes and findings (primary and secondary outcomes measured, main findings and mechanisms of action).

### Quality assessment and synthesis of findings

Trial quality was assessed using five key criteria from the Mixed Methods Appraisal Tool (MMAT) Version 2018 [26]. Papers that did not fully meet the criteria for one or two of the following (blinding, randomisation, comparability of the groups, participant adherence or complete results) were still included, with the limitations noted and considered in the review's overall assessment, as they can still contribute to the context-mechanism-outcome (CMO) configurations.

Narrative analysis [27] began with the descriptive summaries of each of the included studies, allowing reviewers to systematically engage with each study and refine the data extraction elements. To organise the findings, pattern recognition and categorisation were applied. Extracted data were examined for shared characteristics, including: (a) timing of PC initiation (e.g. within 8 weeks of diagnosis, at start of chemotherapy, based on prognosis); (b) model of integration (referral-based, structured collaborative or fully embedded); (c) provider type (nurse-led, doctor-led, multidisciplinary team); (d) intervention components (symptom management, communication and advance care planning) and (e) outcome measures used. Studies were grouped accordingly to enable comparative analysis across these domains.

A quantitative approach was used within each category to describe the frequency of different reported intervention types. Summarising these counts helped identify which interventions were most studied or implemented while also revealing gaps in the evidence base. This process was iterative, involving ongoing refinement of classifications and ensuring coherence across studies. Thematic analysis was conducted using an inductive approach to identify recurring themes and conceptual patterns across included studies. The process followed Braun and Clarke's six-phase framework: (1) familiarisation with extracted data through repeated reading; (2) generation of initial codes manually by two reviewers; (3) searching for themes by grouping related codes; (4) reviewing themes against the original data; (5) defining and naming themes (e.g. symptom management mechanisms, psychological support pathways and system-level integration facilitators) and (6) producing the final synthesis. This approach facilitated the structured synthesis of key insights related to intervention context, core components, implementation facilitators, barriers and reported outcomes, providing a richer understanding of the evidence.

To further contextualise findings, a logic model was developed using the CMO framework [28] aligned with the Donabedian model of health-care quality [29]. The CMO framework was applied to examine how the different contexts and components lead to the outcomes and helped to identify the mechanism of action. To better understand success factors and barriers to implementation, we synthesised evidence from studies reporting both positive and inconclusive findings.

Finally, the transferability of findings to a different resource setting was assessed using the TRANSFER assessment framework [30]. This involved analysis of intervention characteristics (e.g. complexity, resource requirements and feasibility of scale-up); health system factors (e.g. infrastructure, workforce capacity and policy environment); sociocultural considerations (e.g. patient and provider attitudes towards PC and cultural norms surrounding serious illness and end-of-life care). Three independent PC providers from low-resource settings viewed the findings of this systematic review across each intervention characteristic, rating it as causing: no concern, minor concerns, moderate concerns or major concerns, when applied in their settings. Recommendations were extrapolated for the implementation of early/integrated PC delivery in resource-limited settings.

## Results

### Search yield

Electronic database searches yielded  $n = 2,428$  studies. Following deduplication,  $n = 1,523$  articles underwent abstract and title screening,  $n = 175$  full text review, and  $n = 46$  articles were retained for analysis (Figure 1).

### Study characteristics

The retained studies ( $n = 46$ ) were published from 2010 to 2024, and most were ( $n = 40/46$ ) conducted in high-income countries (HICs), with a predominance from North America (United States: 15 and Canada: 4) and Europe (Italy: 3, Belgium: 2, Denmark: 2, Norway: 2, Czech: 1, France: 1, Germany: 1, Switzerland: 1 and United Kingdom: 1). This was followed by Asia (China: 3, Japan: 1, India: 2, Singapore: 1 and South Korea: 1), Oceania (Australia: 3) and South America (Brazil: 1). Of these,  $n = 6$  studies were from middle-income countries:  $n = 2$  from India (a lower-middle-income country) and  $n = 4$  from upper-middle-income countries (Brazil and China).

According to the Global Atlas of PC [9], most of these countries ( $n = 38/46$ ) are at 'advanced' stages of PC development, i.e. level 4a or 4b, indicating specialist PC services integrated into the mainstream health system.

### Quality assessment

RCT quality is reported in Table 3. Most studies ( $n = 40/46$ ) fulfilled 4 of 5 criteria of the MMAT Version 2018, with blinding compromised in  $n = 36$  studies, which is an inherent challenge in PC trials where participants and providers cannot be blinded to the intervention.

Only in one study did participants not adhere to the intervention [31], and in only one other study were the results incomplete, as it was prematurely terminated due to patient death or loss to follow-up [32].

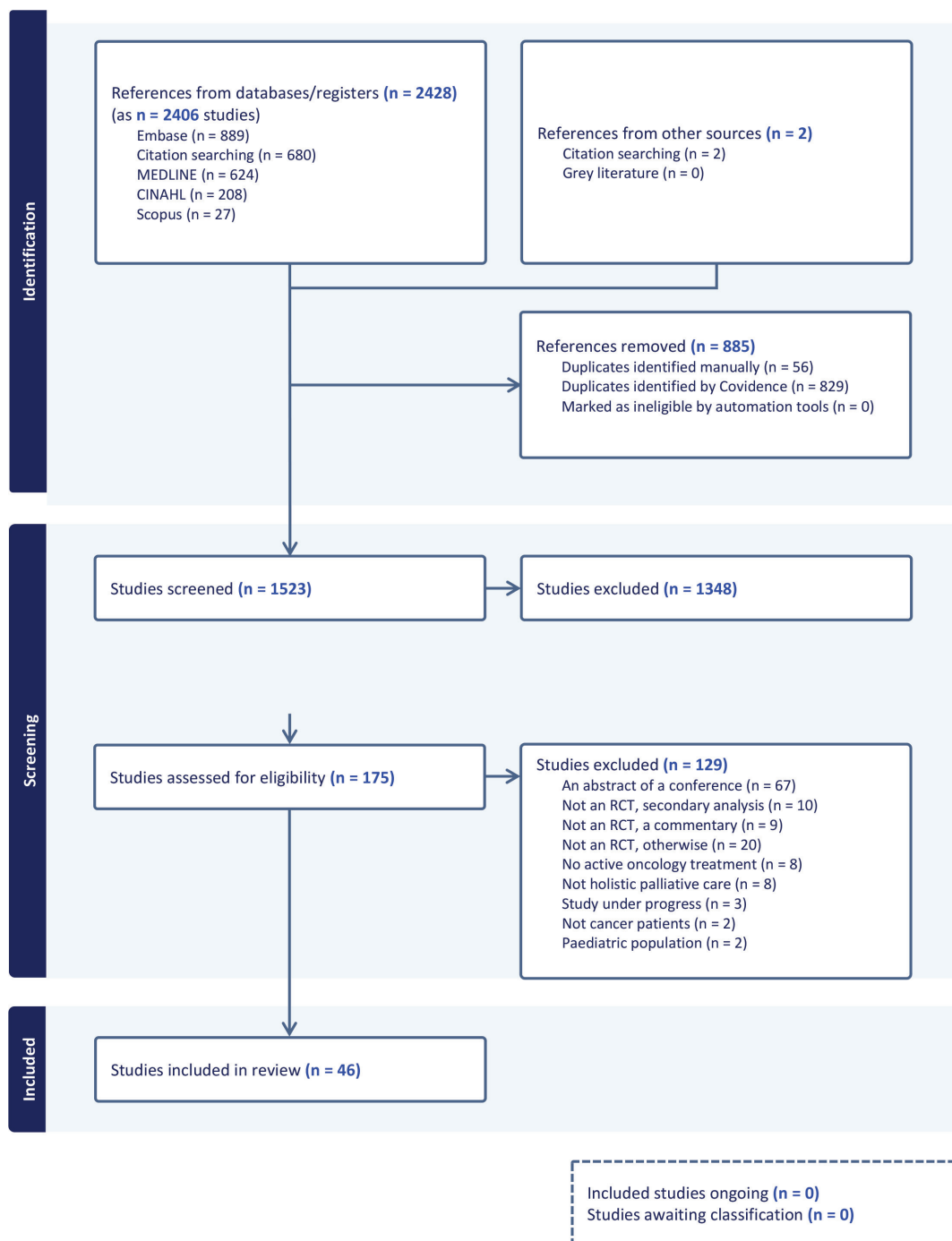


Figure 1. PRISMA flowchart for study selection.

Table 3. Quality assessment results using MMAT tool.

| Study             | Randomisation appropriate | Groups comparable | Data was complete | Assessors blinded | Participants adhered | Study | Randomisation appropriate | Groups comparable | Data was complete | Assessors blinded | Participants adhered | Study            |
|-------------------|---------------------------|-------------------|-------------------|-------------------|----------------------|-------|---------------------------|-------------------|-------------------|-------------------|----------------------|------------------|
| Bakitas, 2015     | ψ                         | ψ                 | ψ                 | ψ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    | McCorkle, 2015   |
| Brims, 2019       | ψ                         | ψ                 | ψ                 | o                 | ψ                    |       | ψ                         | ψ                 | ψ                 | o                 | ψ                    | McDonald, 2017   |
| Chen, 2022        | ψ                         | ψ                 | ψ                 | o                 | o                    |       | ψ                         | ψ                 | ψ                 | ψ                 | ψ                    | Nottelmann 2021  |
| Chvetzoff, 2023   | ψ                         | ψ                 | o                 | o                 | ψ                    |       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Patel, 2022      |
| Collins, 2022     | ψ                         | ψ                 | ψ                 | o                 | ψ                    |       | ψ                         | ψ                 | ψ                 | o                 | ψ                    | Patil, 2021      |
| Datta, 2024       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | o                 | ψ                    | Rodin, 2020      |
| Dionne-Odom, 2016 | ψ                         | ψ                 | ψ                 | ψ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | o                 | ψ                    | Scarpi, 2019     |
| Dionne-Odom, 2022 | ψ                         | ψ                 | ψ                 | ψ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Schenker, 2018   |
| do Carmo, 2017    | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | ψ                 | ψ                    | Schenker, 2021   |
| El-Jawahri, 2017  | ψ                         | ψ                 | ψ                 | ψ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Slama, 2019      |
| El-Jawahri, 2019  | ψ                         | ψ                 | ψ                 | o                 | ψ                    |       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Tattarsall, 2014 |
| El-Jawahri, 2021  | ψ                         | ψ                 | ψ                 | o                 | ψ                    |       | ψ                         | ψ                 | ψ                 | o                 | ψ                    | Temel, 2010      |
| Eychmuller 2021   | ψ                         | ψ                 | ψ                 | ψ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Temel, 2017      |
| Franciosi, 2019   | ψ                         | ψ                 | ψ                 | ψ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | o                 | ψ                    | Temel, 2020      |
| Ferrel, 2021      | ψ                         | ψ                 | ψ                 | o                 | ψ                    |       | Ξ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Treasure, 2020   |
| Greer, 2022       | ψ                         | ψ                 | ψ                 | ψ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Ullrich, 2021    |
| Groenvold, 2017   | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | o                 | ψ                    | Vanbutsele, 2018 |
| Hjermstad, 2021   | Ξ                         | Ξ                 | ψ                 | o                 | ψ                    |       | Ξ                         | ψ                 | ψ                 | o                 | ψ                    | Vanbutsele, 2019 |
| Hjermstad, 2024   | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | o                 | ψ                    | Verma, 2023      |
| Liu, 2023         | ψ                         | ψ                 | ψ                 | o                 | ψ                    |       | Ξ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Woo, 2019        |
| Lu, 2021          | ψ                         | ψ                 | ψ                 | o                 | ψ                    |       | ψ                         | ψ                 | ψ                 | o                 | ψ                    | Yang, 2021       |
| Maltoni, 2016     | ψ                         | ψ                 | ψ                 | o                 | ψ                    |       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Zimmermann 2014  |
| Matsumoto 2022    | Ξ                         | ψ                 | ψ                 | ψ                 | ψ                    |       | ψ                         | ψ                 | ψ                 | Ξ                 | ψ                    | Zimmermann 2022  |

ψ = yes (green), o = no (pink), Ξ = information unavailable (blue)



Major weaknesses identified across the included studies include: (a) lack of blinding in 78% of studies, potentially introducing performance and detection bias; (b) short follow-up periods (only seven studies followed patients beyond 12 months), limiting assessment of long-term outcomes; (c) predominant focus on solid tumours (42/46 studies), with haematological malignancies underrepresented; (d) limited reporting of implementation fidelity and intervention adherence and (e) potential selection bias, as trial participants may not represent the broader oncology population. These limitations are considered in the review's overall assessment while retaining studies for their contribution to CMO configurations'.

## Setting and population

### Setting

Studies were conducted in comprehensive or academic cancer centres ( $n = 19$ ), in general hospitals with oncology departments ( $n = 5$ ) or community-based facilities ( $n = 4$ ) or in a combination of such facilities ( $n = 18$ ). The majority of studies (22/46) were conducted in an outpatient setting [31–52]. Six studies were conducted in an inpatient setting [53–58] or both inpatient and outpatient setting ( $n = 18$ ) [59–75].

### Population

The majority of cancer types were solid tumours (42/46). Three studies dealt with haematologic malignancies [54, 65, 69], and one study included both solid and haematologic malignancies [60]. Regarding disease stage at study entry, most studies (37/46) indicated that the disease was in an incurable, refractory, metastatic or advanced stage. Six studies specifically described their population as newly diagnosed with cancer [46, 47, 52, 59, 70, 71].

### Early PC

Thirty-three studies were explicitly labeled as 'early': [31–34, 36, 38, 39, 41, 42, 45–47, 49–54, 56, 59–64, 68–78]. 'Early' was described as the initiation of PC within a narrow timeframe following a diagnosis of advanced, metastatic or incurable cancer. This criterion was applied in  $n = 16$  studies, which specified windows ranging from 30 to 60 days, with 'within 8 weeks of diagnosis' serving as the most frequent benchmark [33, 34, 39, 41, 42, 45, 46, 49, 50, 59, 60, 63, 64, 71, 73, 76]. A second, smaller group of studies defined 'early' by anchoring it to specific clinical events or treatment phases rather than the date of diagnosis. This heterogeneous group encompassed several distinct approaches. Some defined 'early' relative to the start of a defined line of anticancer therapy, such as first-line chemotherapy [56, 77], the last anticancer line [38], at the beginning of palliative systemic therapy [51] or at a specific treatment decision-point, such as when deciding on third or fourth line chemotherapy [32] or palliative chemotherapy [68] or during hospitalisation for intensive treatment [54, 69]. Others used prognostic criteria, intervening for patients with a life expectancy greater than 6 months [75, 78]. Finally, a few studies employed systematic or trigger-based models, initiating care 'soon after randomisation' [62], after screening for palliative needs [36], at 'evidence-based disease-specific trigger points' [61] or at first contact post-diagnosis [52].

## The interventions

### Integrated PC

Thirty-one studies explicitly named or described their PC model as 'integrated' in their title or abstract [33, 37, 40–44, 47, 49, 51, 53, 54–56, 58–67, 69–73, 76, 77].

'Integrated' consistently signalled a structured, collaborative approach where PC was deliberately combined with standard oncology care from a defined starting point. The meaning of 'integration' most frequently involved co-located, concurrent care delivery [47, 70, 71], where the PC team worked alongside the oncology team in outpatient settings. A core operational feature was interdisciplinary collaboration and shared decision-making, exemplified by regular 'combined clinics' between the PC team and oncologists [32] or co-management between clinicians [59]. Several models described tighter structural unity through shared systems, such as palliative care nurse participation in oncology

meetings and shared records [72], a 'multidisciplinary collaborative team' model [55] or a fully embedded 'co-rounding model' within the oncology team [58]. In other cases, integration was achieved by embedding PC functions into routine oncology practice, such as through oncology nurses delivering primary PC within the same clinic [44] or by incorporating specialist PC as a standard offer within routine cancer care pathways [49].

## Models of PC integration

Of the 46 studies reviewed, the descriptions of the intervention models reveal distinct approaches to how palliative care is organised and delivered. These can be characterised by their core structural principle, the mechanism of interaction between PC and oncology and the degree of their operational integration.

First, a **standalone referral-based consultation** model was employed in several studies, where early PC was the focus, but structural integration was minimal. In this model, the PC team functions as a separate, specialised service. Integration is limited to a formal referral process and subsequent consultation. Interaction is primarily unidirectional; the PC specialist sees the patient and provides recommendations back to the primary oncologist, often via documentation in shared notes or a discharge summary. Studies such as [75, 78] and [52] exemplify this approach, where 'early' is defined by prognosis or time of diagnosis, but the care is delivered in a parallel, consultative fashion. Similarly, studies such as [50] and [36] relied on symptom screening in oncology clinics to trigger a referral to a separate PC service, maintaining distinct care streams. Other studies employing a predominantly referral-based model have been reported [34, 45, 48].

In contrast, the **structured collaborative** model represents a deliberate move towards operational integration while maintaining distinct team identities. This model is defined by scheduled, systematic mechanisms that ensure regular interaction and shared decision making. This includes weekly or regular multidisciplinary team meetings, where PC and oncology clinicians jointly discuss patients [32, 40, 72]. The collaboration extends to co-managing care through a shared plan, which may involve structured, protocol-driven symptom screening with embedded referral pathways [38, 69]. This model [47, 59, 70], effectively blends 'early' initiation (e.g. within 8 weeks of diagnosis) with 'integrated' procedures, ensuring PC is a routine, scheduled component of care rather than an ad hoc consultation. This was the most common model [31, 33, 37, 39, 41–43, 46, 49, 51, 53, 54, 56, 60–64, 67, 77].

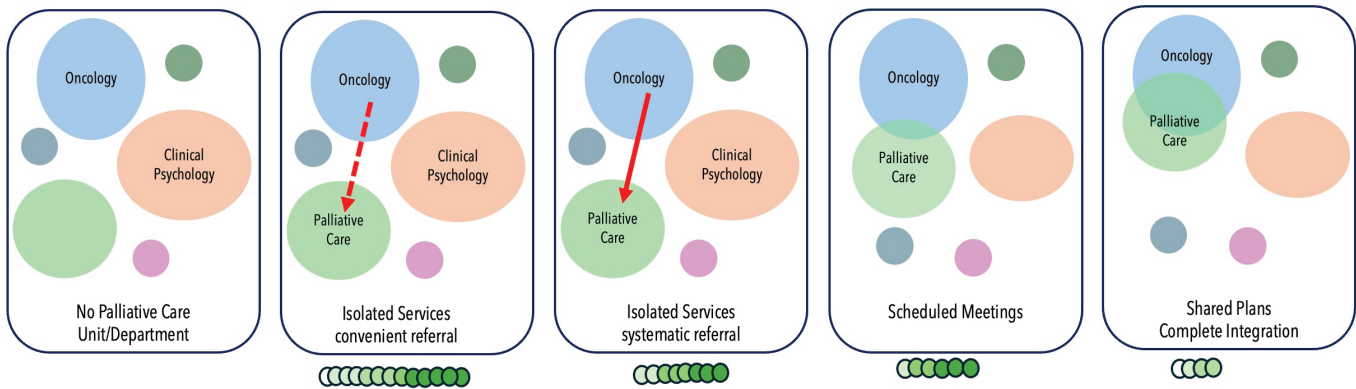
The most fully unified approach is the **embedded or unified care** model, where PC is fully absorbed into the standard oncology care structure. Here, the distinction between 'palliative' and 'oncologic' care blurs, as PC functions are delivered by the core oncology team itself. This is achieved either by training and task-shifting to oncology nurses to provide primary PC within the same clinic visit [44] or through a fully integrated 'co-rounding' team where a PC clinician is a permanent member of the oncology unit, participating in all rounds and decisions [58]. In models such as [55] or the systematic integration in [73], PC is not a separate service to be referred to but is a foundational element of the standard care pathway from the start. This represents the highest degree of integration, often implemented 'early' by design, as the model itself is activated at diagnosis or treatment initiation. Studies with strong elements of this model [65, 66, 76] provide supportive care functions deeply woven into a specific treatment trajectory or clinical trial structure.

Therefore, the spectrum of models ranges from parallel (consultative, referral-based) to collaborative (scheduled co-management) to unified (embedded within oncology). The 'early' aspect is frequently a trigger (time from diagnosis) applied within any of these models, but the depth of 'integration' determines the consistency and quality of the interdisciplinary partnership.

Across these models, the use of systematic symptom screening to trigger referrals ensures that PC is directed to patients with the highest needs, moving beyond ad hoc clinician judgment towards a more equitable and needs-based approach to integration.

Figure 2 shows the progression in interaction between a PC clinic alongside an oncology clinic.

Four additional facilitators were identified as supporting PC integration. Firstly, training was highlighted as a critical facilitator, encompassing training of oncology teams in PC; PC teams in oncology; other health professionals in PC, and carers and patients (psychoeducation) [40, 50, 60, 66, 67, 72].



**Figure 2.** The four images show progression in interaction between a PC clinic alongside an oncology clinic. The circles within the box represent different departments/units within centres/hospitals; the row of small circles below the box each represents an individual study from the review containing that integration model; the darker the green, the more positive/significant are the results in the RCT.

Secondly, the use of specific guidelines and protocols emerged as a key facilitator. The use of specific PC guidelines and protocols, referral guidelines, checklists and bereavement support tools was identified as crucial for structured integration [32, 34, 36, 37, 39, 45, 47, 50, 55, 59, 61, 63, 79]. Thirdly, shared documentation and communication systems facilitated seamless information exchange, including co-located clinics with shared electronic health records [47, 70, 71], PC nurse participation in oncology team meetings [72] and regular interdisciplinary case conferences [32, 40]. Fourthly, institutional leadership and resource allocation, such as dedicated funding for PC positions, protected time for interdisciplinary meetings and visible institutional commitment to integration, emerged as foundational prerequisites [40, 58, 72].

### Structural prerequisites for successful integration

Drawing on the Donabedian framework's structure component, the following structural elements were identified as important for integration to work well:

**Workforce composition and expertise:** Studies with dedicated PC specialists (board-certified physicians, advanced practice nurses with  $\geq 5$  years PC experience) achieved greater improvements in complex symptom management and advance care planning outcomes [31, 32, 37, 41, 47, 50, 52, 54, 58, 59, 61, 63, 64, 66, 70, 71, 74, 75]. However, nurse-led models achieved comparable outcomes for basic symptom management and psychosocial support in adequately trained nurses [34, 44, 60, 72, 73].

**Physical infrastructure:** Co-located clinic space enabling same-day access to both oncology and PC services [47, 70, 71], private consultation rooms for sensitive conversations about prognosis and end-of-life preferences [57] and reliable telephone/telehealth systems for follow-up [45, 60, 68, 75, 78] emerged as important structural enablers.

**Information systems:** Shared electronic medical records accessible to both oncology and PC teams [40, 72], embedded clinical decision support tools (e.g. automated referral triggers based on symptom scores) and systematic documentation templates for advance care planning were associated with better care coordination.

**Governance and accountability structures:** Formal integration agreements defining roles, responsibilities and communication protocols; dedicated leadership with accountability for integration metrics and mechanisms for monitoring quality indicators (e.g. time from diagnosis to PC referral, proportion of patients with documented goals of care) distinguished higher performing integration models [32, 36, 40, 58, 72].

## Intervention components

The majority of studies emphasised the importance of pain and symptom management, communication and decision making, holistic care and family support. Other specific interventions included counselling sessions: [34, 45, 49, 55, 57, 59, 62, 63, 65, 68, 69]; rehabilitation [42]; advanced care planning [43]; inpatient visits [54, 75, 78]; scheduled outpatient PC visits alongside oncology care [31, 37, 39–41, 44–47, 63, 64, 68, 70, 71, 74, 75, 78]; 24-h on-call service [60, 68, 75, 78] and email follow-up [45].

## Providers of the intervention

PC services were delivered mainly by a dyad of doctor and nurse ( $N = 18/46$  studies, 39%), multidisciplinary teams ( $n = 13$ , 28%), a nurse only ( $n = 6$ , 13%), a doctor only ( $n = 5$ , 11%) or another professional ( $n = 4$ , 8%).

First, the doctor–nurse dyad: They are usually highly trained or qualified specialists in PC [31, 32, 37, 41, 47, 50, 52, 54, 58, 59, 61, 63, 64, 66, 70, 71, 74, 75]. Such training ranged from board-certified physicians with 3–5 years of specialist experience, advanced practice nurses with 10–15 years in the field, research nurses trained in symptom management and depression and participants in targeted training sessions.

Second, multidisciplinary teams mainly consisting of physicians, nurses, physiotherapists, psychologists, occupational therapists, dieticians, social workers and chaplains [36, 42, 51, 67]. For some, the team included medical oncologists and specialised oncology nurses [40, 53, 56]. In one, it included oncologists and psycho-oncologists and communication specialists [49]. Further studies mentioned specialists in tumours, nutrition and rehabilitation [55] or advanced practice nurses and mental health clinical nurse specialists [48]. Social workers were emphasised in one study [69]. Two mentioned PC specialists without further details [33, 39].

Third, a sole practitioner or teams, each of a single professional category. The most common profession in this category was nurses [34, 44, 60, 72, 73, 78] (infusion room nurses) [57], followed by doctors [38, 39, 45, 46, 68], psychologists [62, 65], community health workers [43] and navigators [35].

## Outcomes measured

The main constructs measured were quality of life, emotional well-being, survival and healthcare utilisation (Table 4).

## Follow-up period

The total period of assessment largely ranged from 3 to 6 months ( $n = 26/46$  studies). Specifically, the follow-up periods were 2 months: [36, 65]; 3 months: [32, 34, 42, 66, 68, 69, 74]; 4 months: [33, 41, 44, 45, 58, 75]; 6 months: [31, 35, 37, 49, 51, 53, 54, 62, 64, 71, 78]; 12 months: [43, 50, 60]; 20 months: [73]; 24 months: [59, 63, 70]; until death (flexible period): [39, 46, 47, 52, 55] and at irregular time periods: [38, 40, 48, 56, 57, 61, 67, 72].

## Mechanisms of action

These mechanisms are multifactorial and overlapping. They are reported as described in the included studies.

## Physical symptom control and functionality

The effects of early or integrated PC were associated with better control of pain, fatigue, dyspnoea and other distressing symptoms [40]; integrating physiotherapy, occupational therapy and nutritional support to maintain independence and reduce symptom burden [42]; adapting care to cultural, spiritual and personal values, thereby potentially ensuring a holistic approach aligned with patient preferences [35, 37, 42, 65] and regular assessment enabling timely palliative interventions, reducing crisis situations and improving overall well-being [40].

## Psychological and emotional well-being

Interventions were associated with positive effects on patients' psychological health including enhancing coping mechanisms and reducing emotional distress; establishing strong rapport with patients; creating a safe space for discussing fears, anxieties and future planning [64]; guiding them through stress management techniques, emotional resilience training and goal setting; helping them navigate the uncertainty of illness [35, 54, 64, 65]; assisting patients and caregivers and directing them through their care; offering navigation, problem-solving and ensuring they have the necessary psychological resources to handle treatment-related stress [35].

Table 4. Primary and secondary outcome measures used for the studies and identification of studies as early PC, integrated PC or both.

| Study                             | Early or integrated PC | QUALITY OF LIFE | Emotional well-being | Survival | Healthcare utilization |
|-----------------------------------|------------------------|-----------------|----------------------|----------|------------------------|
| Bakitas <i>et al</i> [21, 60]     | Both                   | X               |                      | X        | X                      |
| Brims <i>et al</i> [31]           | Early                  | X               | X                    | X        |                        |
| Chen <i>et al</i> [53]            | Both                   | X               |                      |          |                        |
| Chvetzoff <i>et al</i> [32]       | Early                  | X               |                      |          | X                      |
| Collins <i>et al</i> [61]         | Both                   |                 |                      |          |                        |
| Datta <i>et al</i> [33]           | Both                   | X               |                      |          |                        |
| Dionne-Odom <i>et al</i> [34, 79] | Early                  |                 | X                    |          |                        |
| Dionne-Odom <i>et al</i> [35, 76] | Both                   |                 | X                    |          |                        |
| do Carmo <i>et al</i> [62]        | Both                   | X               | X                    |          |                        |
| El-Jawahri <i>et al</i> [64]      | Both                   |                 |                      |          |                        |
| El-Jawahri <i>et al</i> [65]      | Integrated             | X               | X                    |          |                        |
| El-Jawahri <i>et al</i> [54]      | Both                   | X               | X                    |          |                        |
| Eychmuller <i>et al</i> [63]      | Both                   | X               |                      | X        |                        |
| Ferrell <i>et al</i> [66]         | Integrated             | X               | X                    | X        |                        |
| Franciosi <i>et al</i> [59]       | Both                   | X               | X                    | X        |                        |
| Greer <i>et al</i> [37]           | Integrated             | X               | X                    |          |                        |
| Groenvold <i>et al</i> [36]       | Early                  |                 |                      |          |                        |
| Hjermstad <i>et al</i> [67]       | Integrated             | X               |                      |          | X                      |
| Hjermstad <i>et al</i> [38]       | Early                  | X               |                      |          |                        |
| Liu <i>et al</i> [55]             | Integrated             | X               | X                    |          |                        |
| Lu <i>et al</i> [56]              | Both                   | X               | X                    | X        |                        |
| Maltoni <i>et al</i> [39]         | Early                  | X               |                      |          |                        |
| Matsumoto <i>et al</i> [77]       | Both                   | X               | X                    |          |                        |
| McCorkle <i>et al</i> [40]        | Integrated             | X               | X                    |          |                        |
| McDonald <i>et al</i> [41]        | Both                   | X               | X                    |          |                        |
| Nottelmann <i>et al</i> [42]      | Both                   | X               | X                    |          |                        |
| Patel <i>et al</i> [43]           | Integrated             | X               | X                    | X        | X                      |
| Patil <i>et al</i> [68]           | Early                  | X               | X                    | X        |                        |
| Rodin <i>et al</i> [69]           | Both                   | X               | X                    |          |                        |
| Scarpi <i>et al</i> [46]          | Early                  | X               |                      |          |                        |
| Schenker <i>et al</i> [45]        | Early                  |                 |                      |          |                        |
| Schenker <i>et al</i> [44]        | Integrated             | X               | X                    |          |                        |
| Slama <i>et al</i> [51]           | Both                   | X               | X                    |          |                        |
| Tattersall <i>et al</i> [52]      | Early                  | X               |                      |          |                        |

Continued

**Table 4. Primary and secondary outcome measures used for the studies and identification of studies as early PC, integrated PC or both. Continued**

|                              |            |   |   |   |   |
|------------------------------|------------|---|---|---|---|
| Temel <i>et al</i> [47]      | Both       | X |   | X |   |
| Temel <i>et al</i> [70]      | Both       | X |   |   |   |
| Temel <i>et al</i> [71]      | Both       | X | X |   |   |
| Ullrich <i>et al</i> [49]    | Both       | X |   |   |   |
| Vanbutsele <i>et al</i> [72] | Both       | X | X | X |   |
| Vanbutsele <i>et al</i> [73] | Both       |   |   |   |   |
| Verma <i>et al</i> [74]      | Early      | X | X | X |   |
| Woo <i>et al</i> [50]        | Early      | X |   |   |   |
| Yang <i>et al</i> [58]       | Integrated |   |   |   | X |
| Zimmermann <i>et al</i> [75] | Early      | X | X |   |   |
| Zimmermann <i>et al</i> [78] | Early      | X |   |   |   |

### Optimising end-of-life experiences for patients and families

Interventions were associated with facilitated discussions about patients' values, treatment preferences and goals of care, which may help to ensure that medical decisions align with their wishes including the use of Advanced Care Planning [35, 41, 43, 64]; engaging caregivers early in the process may have helped them understand the disease progression, reducing uncertainty and emotional burden [34]; reducing caregiver distress by improving symptom control and patient comfort; reducing caregiver burnout, depression and complicated grief post-bereavement [34, 79]; prioritising comfort over non-beneficial interventions, which was associated with reduced unnecessary hospitalisations, Intensive Care Unit (ICU) admissions and burdensome treatments at the end of life [38, 63, 66]; patients and families may feel supported, heard and included in decision making, potentially leading to a more meaningful and less traumatic end-of-life experience [31, 41, 43, 57, 61, 75].

### Improving healthcare utilisation and cost-effectiveness

Early and/or integrated PC was associated with efficient healthcare resource use and equitable access by: effective symptom management associated with unnecessary emergency admissions, which may allow patients to remain at home or in less intensive care settings [43, 58, 61]; improving symptom screening and timely interventions, better resource allocation, potentially reducing the financial burden on both healthcare systems and families [43, 46, 74].

### Strengthening interdisciplinary collaboration in oncology care

The integration of PC within oncology was associated with enhanced care coordination and treatment outcomes by: close collaboration between oncology and PC teams associated with timely pain relief and discharge planning, potentially minimising prolonged hospitalisations [32, 36, 58]; regular interdisciplinary meetings helped align treatment goals, avoided unnecessary interventions and ensured smoother transitions between care settings [40]; a structured approach to shared decision making ensures that patients' values guided treatment decisions, promoting dignity and respect in their final phase of life [42, 45].

Here are several points to consider when comparing RCT and real-world practice. Trial patients are usually healthier than real-world populations. Staffing ratios in trials exceed what most systems provide and fee-for-service models generally do not reimburse interdisciplinary meetings or care coordination. Workforce shortages cause waiting times of several weeks for PC, undermining the 'early' principle. Oncologists may resist referral, viewing it as 'giving up', unlike in trial settings. Interventions are often diluted in practice compared to controlled trial settings.

## Transferability

Assessment using the TRANSFER framework (Table 5) found that the review's findings might be transferred to low- and middle-income country [80] settings, although no studies were conducted in low-income countries. Direct application is hindered by differences in the study populations and settings. The evidence base is drawn overwhelmingly from HICs and specialised cancer centres, which differ substantially from the patient demographics, health system structures and resource levels typical in LMICs. Patients in LMICs often present with later-stage disease, have lower health literacy, face greater barriers to healthcare access (transportation, out-of-pocket costs) and may have cultural barriers to PC acceptance [81]. Furthermore, the well-resourced, interdisciplinary models of physicians, nurses, psychologists, social workers and chaplains described contrast with the fragmented oncology and PC services common in many LMICs, where a single nurse may cover multiple roles and where key facilitators like institutional support are weak, and barriers like policy gaps are pronounced [82, 83]. Many LMICs also lack basic components assumed in the included studies: reliable opioid availability, trained PC workforce, PC included in national essential health benefit packages, functional referral systems and electronic health records. Consequently, direct applicability of findings to low-resource settings remains uncertain and largely theoretical. Significant adaptation to the local context is therefore essential.

The core principles of early and integrated PC (timely access, collaboration and symptom management) remain relevant. However, feasible implementation would require modification, including simplifying complex team structures, shifting tasks to nurses or community health workers and using telemedicine to extend specialist support. Creating locally appropriate protocols, rather than importing manuals, is crucial. Foundational requirements, such as reliable access to essential medications like opioids, private spaces for consultations and basic systems for patient follow-up, must also be secured. While outcomes like quality of life and emotional well-being are universal goals, the tools to measure them may need to be shortened, translated and adapted for populations with lower literacy to ensure that they are practical and meaningful in LMIC clinical practice and research.

Figure 3 presents the process logic model illustrating the hypothesised relationships between the core mechanisms of PC and their resulting outcomes.

## Discussion

Based on this systematic review of 46 studies, the terms 'early' and 'integrated', while often used together, describe related but conceptually distinct dimensions of PC intervention design. The distinction lies in what each term prioritises.

'Early' is fundamentally a temporal construct, concerned with when PC is initiated relative to a clinical milestone. The analysis shows this was most commonly operationalised as a protocol mandating intervention within a defined timeframe, most often within 8 weeks of a diagnosis of advanced, metastatic or incurable cancer [39, 60, 71]. Alternative definitions anchored 'early' to the start of a specific treatment line, such as first-line chemotherapy [56] or to a prognosis-based window of 6–24 months [75]. A specific group is related to screening for PC needs or immediately upon first contact post-diagnosis. In practice, these definitions function as practical triggers to initiate care. The core intention across all models is to ensure PC is provided concurrently with active oncology treatment from near the beginning of the advanced illness trajectory.

In contrast, 'integrated' is an organisational and relational construct. It describes how PC and standard oncology care are structured to work together. Our review identified a spectrum of integration models, from basic co-located concurrent care to deep structural unification. The defining feature was not timing, but the mechanism of collaboration. This ranged from scheduled co-management in shared clinics [47, 70] and regular interdisciplinary meetings [32, 72] to fully embedded models, where PC functions were performed by the oncology team itself, such as through nurse-led primary PC [44] or a 'co-rounding' team structure [58]. Therefore, an intervention can be 'early' without being 'integrated' (e.g. a standalone early referral) and 'integrated' without being strictly 'early' (e.g. a pathway integrated at the start of last-line therapy). Studies that combine both principles – implementing a protocol-driven early timing rule within a structured collaborative care model – appear to be the most effective, positioning PC as a routine, co-managed component of standard oncology practice from diagnosis.

Several facilitators of PC integration were identified in this review, including systematic symptom screening, training and the use of clinical guidelines.

The value of systematic symptom screening and needs assessment as a facilitator is well established, enabling the proactive identification of patients with unmet PC needs rather than relying on ad hoc clinician referral [84, 85].



Table 5. Results of the TRANSER assessment of SR findings into resource limited settings.

| TRANSFER factor                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     | Findings and considerations in a low- and middle- income setting                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    | Grade          |
|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------|
| <b>Population</b><br>Demographic characteristics (ethnicity, socio-economic aspects, age, workforce participation, and education] the participants' illness or condition, comorbidities, and patient acceptability.                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 | <b>Target population</b> <ul style="list-style-type: none"> <li>• Most studies are from HICs, with few from LMICs and UMICs, indicating a strong evidence base on well-resourced healthcare systems and higher levels of literacy in the population. <i>Consider patient and family empowerment through knowledge about the disease, its complications and role of PC.</i></li> <li>• Transferability requires considering the prevalence of specific cancer types and stages at presentation. The majority of the RCT were conducted on patients with solid tumours and some haematological malignancies which would be similar in LMICs.</li> </ul>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               | Minor concerns |
| <b>Intervention</b><br><i>Intervention characteristics: intervention delivery:</i><br>a] whether there is a possibility for tailoring an intervention,<br>b] the intensity and duration of an intervention, the materials/manuals used to deliver an intervention<br>c] the settings in which the intervention is delivered (hospital, home, etc.),<br>d] whether there are standard procedures for the intervention,<br>e] who pays for the intervention, and whether any other general details related to intervention delivery could influence transferability.                                                                                                                                                                                  | <b>Intervention components:</b> <ul style="list-style-type: none"> <li>• The review findings are from settings with advanced stages of PC integration. Local adaptation is needed for settings with preliminary or isolated integration, such as in China, Brazil, and India.</li> <li>• Intervention components such as interdisciplinary teams may need simplification or modification to fit the local healthcare workforce and resources. Cultural considerations, like family involvement in care decisions, should be factored into the adaptation processes.</li> <li>• Resource limited setting might find it difficult to produce written standard procedures or manuals.</li> <li>• <i>Consider infrastructural needs for provision of PC: a] ensuring continuity of care through telephone calls, computers for documentation and system of follow-up; b] ensuring good communication and conducting family meetings through required space which provides privacy; c] ensure access to medications for symptom control, especially opioids, and ensure sufficient dispensed quantities</i></li> </ul>                                   | Minor concerns |
| <b>Implementation</b><br><i>individual service providers:</i> how the number, and type of personnel, and their skills, training, and other characteristics could influence transferability, practices around monitoring and supervision of the service providers, their compliance, and factors that influence the motivation of service providers<br><i>the organization responsible for implementing an intervention:</i><br>a] the number of essential resources that are available to the implementing organization,<br>b] the culture of the implementing organization concerning mission, mandates, climate, how ready the organization is for implementation, and whether it is feasible for the organization to implement the intervention. | <ul style="list-style-type: none"> <li>• Effective transferability would involve assessing the availability and accessibility of PC services locally.</li> <li>• Training local healthcare providers and task shifting can extend specific skills to different cadres, as well as use of technology, e.g. consultation by phone with PC specialists.</li> <li>• Need to set standards of practice and their evaluation e.g. referrals to PC should not be subjective or optional.</li> <li>• <i>Consider the doctor nurse dyad in the provision of PC services, as there is scarcity of trained health professionals in PC in LMIC (i.e. difficulty of having a complete qualified and experienced team]; consider task shifting into nurses and CWH according to availability of staff; consider training of the oncology teams in the basics of PC to provide basic services and assist in early referrals; consider adoption and adaptation of guidelines and tools from similar settings more suitable with reduced workforce, e.g. simplified and shorter tools; consider organizational changes to compensate for PC services.</i></li> </ul> | Minor concerns |

Continued



Table 5. Results of the TRANSER assessment of SR findings into resource limited settings. *Continued*

|                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              |                   |
|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------|
| <b>Comparison</b><br>characteristics of the comparison condition and quality of the usual services to which an interventions effect is being compared.                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              | <b>Comparison:</b> <ul style="list-style-type: none"> <li>• Conducting the interventions in settings where PC is new or not integrated might be more productive. In LMIC it is expected to find minimal generalist PC or basics of PC diffused into medical practice.</li> </ul>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             | No concerns       |
| <b>Outcomes</b><br>how the outcome was measured, length of follow-up, key outcomes, or adverse outcomes.                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            | <b>Outcomes:</b> <ul style="list-style-type: none"> <li>• Follow up advised for not less than 12 weeks. In LMIC we need to modify the tools to make them less lengthy and thus encourage patients' compliance, relevant to more illiteracy levels and IT illiteracy. The studies reported significant improvements in quality of life, emotional well-being, caregiver support, and healthcare utilization.</li> </ul>                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       | No concerns       |
| <b>Environmental context</b><br>temporal context, political context (political acceptability, governing system, etc.); regulatory context, systems context (organization of health or welfare care, employment regulations or practices], social context (social cohesion, levels of community trust, presence of racism], other interventions (an environment where multiple competing interventions are implemented concurrently, or where one intervention is closely related to participation in another intervention], and the geographic or physical setting.<br><b>Researcher's conduct</b><br>how the researcher's conduct may influence transferability: participation rate, how participants were selected, eligibility criteria of participants in a study, length and details of the run-in period and how participants were recruited. | <b>Context:</b> <ul style="list-style-type: none"> <li>• The studies were predominantly conducted in comprehensive cancer centres and oncology clinics, these settings may not be highly standardized in LMICs.</li> <li>• Facilitators: strong institutional support, interdisciplinary teamwork, and structured training programs were key facilitators in the reviewed studies.</li> <li>• Barriers included limited generalizability due to single-centre studies, language barriers, and resource constraints.</li> <li>• Transferring findings would involve leveraging local facilitators such as existing oncology infrastructure, policy support, and community engagement. Addressing barriers might involve policy advocacy for PC integration, culturally sensitive training programs, and increasing awareness about early PC benefits among healthcare providers and patients.</li> <li>• <b>Implementation:</b> <ul style="list-style-type: none"> <li>• Implementing these interventions involves developing protocols, training healthcare teams, and establishing monitoring systems.</li> <li>• Ongoing evaluation and feedback mechanisms are crucial for continuous improvement and adaptation.</li> <li>• Ensure proper conduct of the control arm team to avoid contamination.</li> </ul> </li> </ul> | Moderate concerns |

The importance of training for effective integration is reinforced by systematic reviews [5, 18], which emphasise the importance of education for healthcare providers and patients. A prior review suggests that the use of structured guidelines and protocols can enhance the delivery of PC and ensure that best practices are consistently followed [86]. Furthermore, standardised guidelines improve the consistency and quality of PC, with studies [11, 87] demonstrating the effectiveness of using protocols in the integration of PC.

The interventions in this review demonstrate a consistent, multi-faceted impact on patient and family outcomes, which can be understood through a mechanism-to-effect pathway. The fundamental achievement is the improvement in quality of life and emotional well-being, congruent with some landmark meta-analyses [5]. This is not a singular effect but the result of a synergistic process. The mechanism begins with proactive, systematic symptom management [40, 42], which establishes physical comfort as a foundation. Concurrently, dedicated, ongoing support from the PC team fosters therapeutic relationships that enhance coping skills and mitigate distress by providing a secure forum for discussing prognosis and fears [65, 76]. This dual support (addressing the physical and the psychological) directly translates into the gains measured by patient-reported outcome measures.

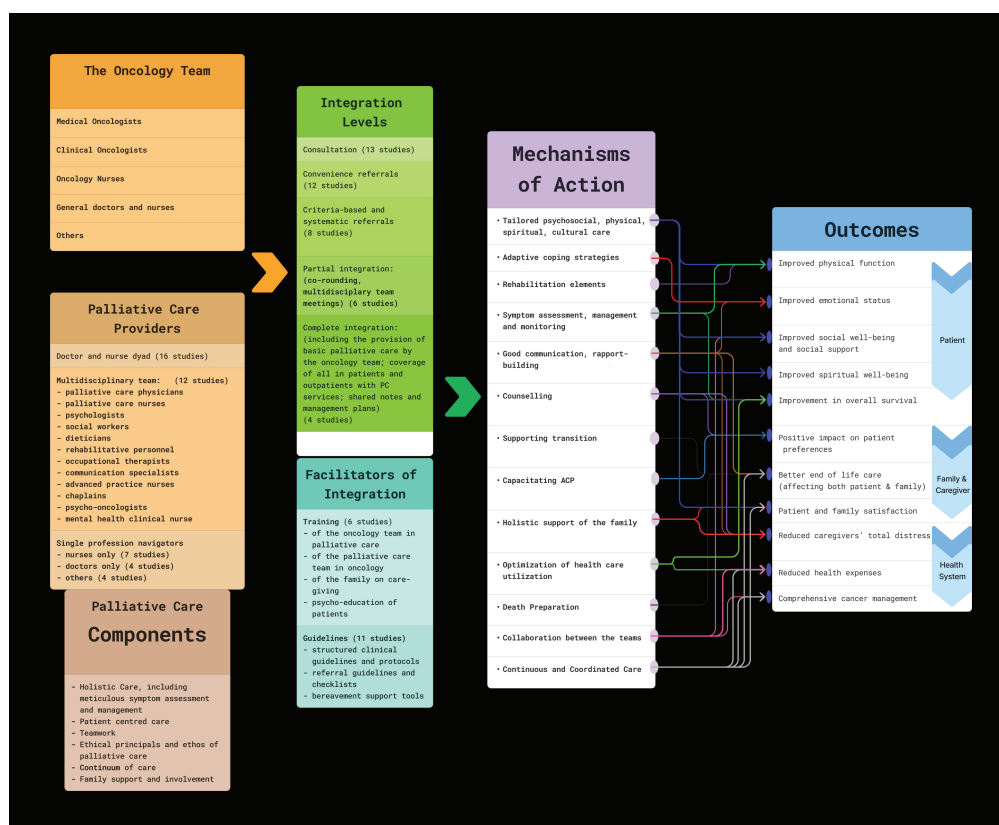
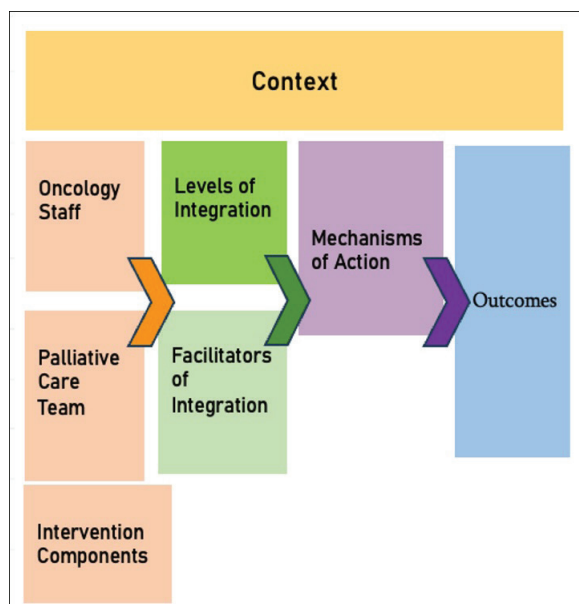


Figure 3. Process logic model for early/integrated PC into oncology. (a) simplified and (b) expanded. The model depicts how these interconnected processes, such as holistic care, patient-centredness and caregiver support, interact to generate positive effects for patients, families and the health system while excluding pathways related solely to healthcare providers.

Beyond quality of life, these models achieve a re-alignment of care with patient values and more appropriate healthcare utilisation. The mechanism here is the structured facilitation of goals-of-care discussions and advance care planning, integrated early in the illness trajectory [41, 43]. This process, supported by ongoing clinician–patient rapport, ensures medical decisions reflect personal preferences, which in turn leads to the observed reductions in aggressive end-of-life treatments and emergency department use [61, 63]. The effect is a care experience that prioritises patient-defined quality over default, often non-beneficial, intensive interventions. Importantly, this value-congruent care also extends support to caregivers, reducing their burden and distress [34].

Finally, the ‘integrated’ dimension of these interventions achieves a system-level effect by enhancing interdisciplinary collaboration. The mechanism is the formalisation of communication channels (through co-location, shared pathways or regular meetings) between oncology and PC teams [32, 72]. This integration streamlines care coordination, enables earlier identification of needs and fosters a unified treatment plan. The outcome is not only more efficient resource use [58], but also the normalisation of palliative principles within standard oncology practice, ensuring a more seamless and supportive continuum of care for all patients facing serious illness. In summary, these interventions work by embedding a parallel process of supportive care that manages symptoms, clarifies goals and coordinates systems, thereby improving patient-centred and health system outcomes.

Most of the studies were conducted in HICs with some level of generalist PC as a part of standard cancer treatment. There, the proven benefits of specialised early and integrated models show the additional value they bring to an existing supportive system. In places where basic palliative support is scarce, the potential benefit could be much greater, but the challenges in delivering these services may also be much larger. Understanding how to adapt and implement these models in resource-limited settings is essential.

This systematic review advances the field by moving beyond confirming the efficacy of ‘early integrated PC as a singular concept. It performs a critical deconstruction, clearly delineating ‘early’ as a temporal/prognostic trigger and ‘integrated’ as an organisational/relational model. By mapping specific intervention components (such as provider structures, i.e. specialist dyads versus multidisciplinary teams; operational mechanisms, e.g. scheduled consultations versus embedded pathways) to these distinct concepts, this review provides a practical framework for implementation. It clarifies that while combined ‘early integrated’ models are potent, effective ‘early’ care can be delivered through simpler, scalable models (e.g. nurse-led protocols) and ‘integration’ can be achieved through task-shifting within oncology teams, rather than only by adding external specialists. This granular analysis of how interventions are built and delivered is a key contribution for translating evidence into heterogeneous clinical settings.

Based on the findings of this review, we propose the following recommendations aimed at translating evidence into meaningful action across clinical, research and policy domains in resource-limited settings.

For clinical practice, we recommend a pragmatic, tiered approach to implementation. Recognising that resources are often constrained, health services should begin by establishing systematic symptom screening for all cancer patients at diagnosis and major treatment junctures. This simple step operationalises the ‘early’ principle by identifying needs proactively. For patients with advanced disease or high symptom burden, a clear, protocol-driven referral pathway to specialist PC (such as referral within 8 weeks of an advanced diagnosis) should be activated, leveraging existing specialist teams or advanced practice nurses. In more resourced settings, such as academic cancer centres, deeper integration can be pursued through multidisciplinary teams or by embedding PC clinicians directly within oncology units. A key, sustainable strategy across all settings is to invest in training oncology nurses and doctors in core PC skills, such as symptom assessment and basic psychosocial support, thereby building integrated capacity from within the existing workforce.

To address critical gaps in the evidence, future research must prioritise pragmatic and context-specific studies. There is an urgent need for studies conducted in LMICs, where the models tested in high-income settings may not be directly applicable. These studies should explore the minimal effective ‘dose’ of PC and determine which personnel models (such as nurse-led versus specialist-led care) are most feasible and effective in different resource environments. We also recommend a full dedicated paper on implementation of early integrated PC in low-resource settings as a separate future direction, including a systematic review of implementation studies and qualitative evidence synthesis from LMICs.

Finally, substantive policy reform is required to create an enabling environment for these models. Policymakers must move beyond rhetorical support and mandate the development of structured clinical pathways that include systematic screening and timed referral criteria for PC,

aligning it with other standards of oncologic care. Policy must also actively support workforce development by funding the training and credentialing of oncology nurses and other allied health professionals in PC competencies and ensure these expanded roles are recognised and reimbursed. Ultimately, financing models must evolve from fee-for-service to value-based systems that incentivise and reward patient-centred outcomes – such as improved quality of life and goal-concordant care – and the reduction of aggressive, non-beneficial treatments at the end of life. This financial imperative is crucial for driving the widespread and sustainable integration of PC into standard oncology practice.

## Conclusion

For LMICs, where patients often present with advanced disease and specialist resources are scarce, the potential impact of integration could be profound, but currently tested models may not be directly feasible. Future efforts must therefore prioritise implementation science in these under-resourced contexts, exploring adapted, sustainable models of care delivery. Ultimately, the goal is to ensure that the holistic support of PC is a seamless part of every cancer patient's journey, irrespective of geography or prognosis.

## Conflicts of interest

The authors declare that they have no competing interests.

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## Author contributions

Nahla Gafer led the study conceptualisation and methodology, performed investigation (screening, review and data extraction) and drafted and revised the manuscript. Nuhamin Gebre contributed to investigation and manuscript review. Khulood Alyami contributed to investigation (data extraction). Matthew Maddocks and Richard Harding contributed to conceptualisation, methodology, validation, review and editing and supervision. All authors approved the final manuscript.

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