End-of-life outcomes with or without early palliative care: a propensity score matched, population-based cancer cohort study

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ABSTRACT

Objectives To investigate whether cancer decedents who received palliative care early (ie, >6 months before death) and not-early had different risk of using hospital care and supportive home care in the last month of life.

Design/setting We identified a population-based cohort of cancer decedents between 2004 and 2014 in Ontario, Canada using linked administrative data. Analysis occurred between August 2017 to March 2019.

Participants We propensity-score matched decedents on receiving early or not-early palliative care using billing claims. We created two groups of matched pairs: one that had Resident Assessment Instrument (RAI) home care assessments in the exposure period (Yes-RAI group) and one that did not (No-RAI group) to control for confounders uniquely available in the assessment, such as health instability and pain. The outcomes were the absolute risk difference between matched pairs in receiving hospital care, supportive home care or hospital death.

Results In the No-RAI group, we identified 36 238 pairs who received early and not-early palliative care. Those in the early palliative care group versus not-early group had a lower absolute risk difference of dying in hospital (−10.0%) and receiving hospital care (−10.4%) and a higher absolute risk difference of receiving supportive home care (23.3%). In the Yes-RAI group, we identified 3586 pairs, where results were similar in magnitude and direction.

Conclusions Cancer decedents who received palliative care earlier than 6 months before death compared with those who did not had a lower absolute risk difference of receiving hospital care and dying in hospital, and an increased absolute risk difference of receiving supportive home care in the last month of life.

INTRODUCTION

Early palliative care is purported to improve quality of life and also avoid unnecessary acute care use, and thus reduce health system costs. Several randomised trials on patients with advanced cancer have shown that early palliative care reduced symptoms and some even had survival benefits.1–3 This evidence led to the oncology clinical practice guideline that supports the early integration of palliative care with standard oncological care.4 5 However, the evidence is mixed as to whether it reduces health services utilisation outcomes at end of life. There are trials that show that resource utilisation at end of life is not different from ‘usual care’.

In particular, many of the trials implemented palliative care interventions close to diagnosis in controlled study settings, which is difficult to implement in the real-world. For example, the US’ Medicare Hospice Benefit requires a physician to certify an expected death within 6 months.6 Additionally, many observational studies have found positive

Strengths and limitations of this study

► This large population-based cohort study of all cancer decedents in Ontario, Canada from 2004 to 2014 uses consistent exposure and outcome definitions over a long period of time, which provides high external validity in real-world settings.

► The study used propensity scores to match decedents who received palliative care earlier than 6 months before death compared with those who did not, thereby reducing selection bias among those who receive early palliative care.

► Our study included and controlled for previously unmeasured confounders known to be associated with receipt of early palliative care (ie, worse pain, Activities of Daily Living (ADL) dependency, depression, cognitive decline and health instability) derived from home care assessment data.

► The study matches those who have similar propensity to have received early palliative care, but this may not represent the entire population of cancer decedents.

► The study does not directly measure patient preferences, which is a confounder for use of early palliative care.

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associations between early palliative care and reduced likelihood to receive aggressive care at end of life (eg, reduced hospitalisations and hospital deaths).\textsuperscript{7–12} However, observational studies are limited by selection bias, namely those who get early palliative care may be different from those who do not (eg, are sicker or more symptomatic in ways that are unmeasured). This is summarised in a large systematic review on early palliative care interventions, which found mixed evidence of benefits and noted key methodological issues of selection bias, as well as large variation in the definitions of when ‘early’ began, the interventions themselves and usual care.\textsuperscript{13} Thus, the evidence that early palliative care reduces late-life acute care use (particularly when it does not begin at diagnosis) is unclear. This gap has important health resource planning and economic implications.

By examining patients with cancer in the universal health system of Ontario, Canada, we are able to address prior limitations, namely standardising definitions for ‘early’ palliative care, usual care and the palliative care intervention. Usual care in Ontario means that patients with cancer have access to publicly subsidised palliative care in the form of: a palliative care outpatient clinic (eg, multidisciplinary pain and symptom management clinic); palliative home care services by a nurse of personal care worker; or a family doctor providing palliative care via clinic or rarely via home visit. Generally, these three services are independent of one another and uncoordinated.\textsuperscript{14} This contrasts the community-based, multidisciplinary team approach of palliative care delivery found in the USA via home hospice care\textsuperscript{6} or in the UK via Macmillan cancer support programme.\textsuperscript{15} Although a small minority of patients might have access to a multidisciplinary, specialist palliative care team that makes home visits or a residential hospice, especially if they lived in a major city, this is haphazard and accessed typically in the last weeks of life.\textsuperscript{16} If the patients were hospitalised, they could also receive a consult from a palliative care doctor individually or a multidisciplinary team (eg, admitted to a palliative care unit) in the hospital. Unfortunately, data show palliative care services are often used very late in the disease trajectory or not at all. For example, in Ontario, Canada, palliative care services are used in 50\% of all deaths for a median of 30 days before death.\textsuperscript{17} In the USA, statistics are very similar, where palliative care via the Medicare Hospice Benefit is used in 45\% of all deaths for a median of 17 days before death.\textsuperscript{18}

Specifically this study investigated the impact of receiving palliative care early (at least 6 months before death) versus not-early on outcomes in the last 30 days of life. We used propensity score matching to reduce selection bias in observational cohorts around receipt of early palliative care.

\textbf{METHODS}

\textbf{Study design and data sources}

We performed a population-based, retrospective cohort study of all cancer decedents in Ontario, Canada from 2004 to 2014. We used propensity score matching to match decedents having received palliative care early (ie, between 12 and 6 months before death) to those who did not (ie, received palliative care late or not at all). We linked administrative databases housed at Institute for Clinical Evaluative Sciences (ICES) (formally known as the ICES) including the: Ontario Cancer Registry (cancer diagnosis), Vital Statistics Registry (death date), Discharge Abstract Database (hospitalisations), National Ambulatory Care Reporting System (emergency department use), physician billings, Statistics Canada (sociodemographic data like income and rurality) and the Home Care Database, which includes all Resident Assessment Instrument-Home Care (RAI-HC) assessments. Datasets were linked using unique encoded identifiers and analysed at ICES.

\textbf{Study population}

We included decedents who had a cancer diagnosis in the Ontario Cancer Registry and a death caused by cancer as per the provincial Vital Statistics registry. Those whose cancer diagnosis was 6 months or less from death were excluded as they were not eligible for the exposure.

\textbf{Exposure}

In the exposure period (ie, between 12 and 6 months before death),\textsuperscript{19} access to early palliative care was defined as having received: homecare with an palliative care intent; a physician consult for palliative care in an inpatient admission (including complex continuing care), outpatient clinic or a home visit setting; or a hospitalisation where palliative care was listed as the main reason for admission, as per prior research.\textsuperscript{17} Once a patient was identified as having received early palliative care, they remained in the exposed group for analysis.

\textbf{Outcomes}

Outcomes were death in an acute care bed, and the aggregate measures of aggressive care and supportive home care in the last 30 days of life respectively. Aggressive care was defined as one or a combination of ≥1 emergency department visit, hospital admission or intensive care unit (ICU) admission.\textsuperscript{20} Supportive home care was defined as one or a combination of physician house call for palliative care, end-of-life homecare nursing or end-of-life personal support at home.\textsuperscript{21} Each outcome was handled as a binary variable (yes/no).

\textbf{Statistical analysis}

To reduce selection bias for decedents who were exposed to early palliative care, we used propensity score matching to create a similar comparison group of unexposed decedents (not-early). The propensity score is an individual’s probability of receiving early palliative care, given the values of their baseline measured covariates. Matching on the propensity score can estimate the effect of the intervention, which is unbiased by differences in the distributions of measured baseline covariates.\textsuperscript{22 23} Our methods matches two individuals who have the same propensity...
to receive early palliative care in the exposure period, though one got early palliative care and one did not.

A priori we decided to examine the group who received long-stay home care services (ie, expected to receive at least 60 days of home care) and thus had an RAI-HC assessment in the exposure period separately. Of note, long-stay home care patients either received standard homecare (unexposed) or palliative homecare (exposed) services. This allowed us to control for additional confounders associated with receipt of early palliative care that are uniquely available in the RAI-HC. Therefore, we created two mutually exclusive groups of matched pairs and each pair consists of an exposed and unexposed decedent. One group is called the No-RAI group; the other the Yes-RAI group.

For the No-RAI group, all pairs were hard matched before the exposure period on: age at death, sex, cancer type, cancer stage (where available) and the logit of the propensity score (callipers of width less than or equal to 0.2 of the SD of the logit of the propensity score). We estimated the propensity score using a logistic regression model with exposure to early palliative care as the independent variable. The predictor variables in the propensity score regression included: income quintile, rurality, health region, prior hospital utilisation in months 24–12 before death, Deyo-modified Charlson Comorbidity Score in months 24–12 before death, index year of death, and having had radiation or cancer surgery.

For the Yes-RAI group, we use additional data from the RAI-HC, which is a standardised assessment for all long-stay home care patients in Ontario, corresponding to the Minimum Data Set in the USA. In addition to matching procedure noted above for the No-RAI group, pairs were hard matched on health instability using the Changes in Health, End-stage Disease, Signs and Symptoms (CHESS) scale. The following items were also included in the propensity score regression: functional performance and dependency using the Activities of Daily Living (ADL) Self-performance Hierarchy Scale; depression using the Depression Rating Scale; cognitive impairment using the Cognitive Performance Scale; pain intensity using the Pain Scale and living with a primary or secondary caregiver.

Because the distributions of covariates were well balanced and not statistically different after matching the exposed and unexposed patients in both the No-RAI and Yes-RAI groups, we did not need to employ any regression methods for examining the exposure-outcomes relationship; thus for each outcome, we determined the absolute risk difference between the matched exposed and unexposed individuals in both Yes-RAI and No-RAI groups. We used McNemar’s test to determine statistical significance of the estimated risk difference. Differences in risk between the exposure and control groups for each outcome were assessed using standardised differences. Standardised differences are more appropriate to use in this population-based study as they are not influenced by sample size (unlike p values). Analysis was performed using SAS Enterprise Guide, V.7.1 (SAS Institute).
As a sensitivity analysis, we divided the not-early group—that is, unexposed group—into late palliative care (ie, only received palliative care in the last 6 months of life) and never received palliative care. We conducted a sensitivity analysis to examine early versus late and early versus never subgroups separately in an attempt to control for unmeasured patient preferences. The hypotheses were that some patients may refuse palliative care altogether (which would appear in our data as never receiving any palliative care services even near death); and other patients might have been willing to receive palliative care but were offered it late (which would appear in our data as receiving it in the final 6 months of life). Analysing the late users to the early users specifically was an attempt to separate out those patients who might have refused palliative care as per their preference. The study is reported using the Strengthening the Reporting of Observational Studies in Epidemiology framework for observational studies.

Patient and public involvement
Patients and the public were not involved in this research.

RESULTS
Patients
After excluding those with a cancer diagnosis within 6 months of death (n=84673), our overall eligible cohort consisted of 144306 cancer decedents in Ontario between 2004 and 2014, of which 53959 (37.4%) received early palliative care 12–6 months before death. Eighty-nine percent (n=128248) of the overall cohort did not have an RAHC in the exposure period (No-RAI) and they were matched separately than the 11% (n=16058) who did have the assessment (Yes-RAI) (figure 1). Baseline characteristics before propensity score matching are shown in online supplemental appendix S1, and those after propensity score matching are shown in table 1.

In the No-RAI group, we matched 82.6% of patients who received early palliative care for a total of 36238 matched pairs. After matching, the decedent covariate distributions were nearly identical between the two groups. For instance, average age was 69, 23.5% had lung cancer and 14.2% had stage IV disease. In the No-RAI group, during the exposure period the group received 53787 palliative care services, of which approximately 40% of services were homecare and 40% were outpatient physician billings. The first initiation of early palliative care was about 330 days before death. In the last 6 months of life, the early group received 8448 palliative care services (same distributions as No-RAI group) whereas the late group received 4664 palliative care services (16% homecare, 38% physician consults and 37% hospital admissions).

Aggressive care
Among matched pairs in the No-RAI group, those who received early palliative care had lower risk difference of the aggressive care outcomes compared with the not-early group (table 2). 38.1% of the early palliative care decedents died in hospital, compared with 48.1% of the non-early palliative care group, resulting in a lower absolute risk difference of 10.0%. Similarly, the aggregate measure of aggressive care was lower by 10.4% among early palliative care decedents. The early palliative care decedents have a lower absolute risk difference of an emergency department visit (9.7%), hospital admission (10.1%) and ICU admission (4.4%) in the last month of life compared with the not-early group.

Among matched pairs in the Yes-RAI groups, we found similar results in the direction and magnitude of the absolute risk differences favouring early palliative care. Note, McNemar’s tests for matched pairs were significant (p<0.0001 for all measures). Further, the sensitivity analyses in the Yes-RAI and No-RAI groups separately, looking at matched pairs of early vs late palliative care and early versus never palliative care, respectively, showed that the early palliative care group consistently had lower absolute risk differences for all outcomes, in similar magnitudes (online supplemental appendix S2).

Supportive home care
Among the matched pairs in the No-RAI group, those who received early palliative care had higher risk of receiving supportive home care outcomes compared with the not-early group. (table 3) The aggregate measure of supportive home care was higher by 23% among early palliative care decedents vs not-early decedents. 56.2% of the early palliative care decedents had any end-of-life home care nursing in the last 30 days, compared with 34.0% of the non-early palliative care group, resulting in a lower absolute risk difference of 22.2%. The early palliative care decedents have a higher absolute risk of having a physician house call (10.2%) and an end-of-life personal support worker in the last month of life (16.0%) versus not-early decedents.

Among the matched pairs in the Yes-RAI groups, we found similar trends in direction, but at larger
| Variables that were hard matched                        | No-RAI | Early palliative care (N=36238) | Yes-RAI | Early palliative care (N=3568) |
|--------------------------------------------------------|--------|---------------------------------|---------|-----------------------------|
| Mean age ±SD                                           | 69.43±12.84 | 69.36±12.87                     | 76.63±11.25 | 76.46±11.19                  |
| Cancer type at diagnosis                               |        |                                 |         |                             |
| Breast                                                 | 4126 (11.4) | 4126 (11.4)                     | 433 (12.1) | 433 (12.1)                  |
| Colorectal                                             | 5266 (14.5) | 5266 (14.5)                     | 722 (20.1) | 722 (20.1)                  |
| Haematology                                            | 2982 (8.2) | 2982 (8.2)                      | 479 (13.4) | 479 (13.4)                  |
| Lung                                                   | 8530 (23.5) | 8530 (23.5)                     | 548 (15.3) | 548 (15.3)                  |
| Prostate                                               | 3053 (8.4) | 3053 (8.4)                      | 486 (13.6) | 486 (13.6)                  |
| Stage at diagnosis                                     |        |                                 |         |                             |
| Stage III                                              | 3726 (10.3) | 3726 (10.3)                     | 275 (7.7)  | 275 (7.7)                   |
| Stage IV                                               | 5151 (14.2) | 5151 (14.2)                     | 329 (9.2)  | 329 (9.2)                   |
| Unavailable                                            | 24631 (68.0) | 24631 (68.0)                   | 2749 (76.7) | 2749 (76.7)                  |
| CHESS Score (when RAI-HC completed)                    |        |                                 |         |                             |
| No health instability                                  | –      | –                               | 921 (25.7) | 921 (25.7)                  |
| Low health instability                                 | –      | –                               | 2242 (62.5) | 2242 (62.5)                 |
| Moderate health instability                            | –      | –                               | 380 (10.6) | 380 (10.6)                  |
| Severe health instability                              | –      | –                               | 43 (1.2)   | 43 (1.2)                    |
| Variables within the Propensity Score                  |        |                                 |         |                             |
| Lowest income quintile                                 | 7058 (19.5) | 7146 (19.7)                     | 776 (21.6) | 790 (22.0)                  |
| Highest income quintile                                | 7102 (19.6) | 7130 (19.7)                     | 626 (17.5) | 622 (17.3)                  |
| Lives in Rural Community                               | 5206 (14.4) | 5236 (14.4)                     | 579 (16.1) | 568 (15.8)                  |
| Deyo-Charlson Comorbidity Score (≥1)                   | 12026 (33.2) | 12540 (34.6)                   | 1483 (41.4) | 1426 (39.8)                 |
| Had Radiation since diagnosis                          | 22337 (61.6) | 21982 (60.7)                    | 1894 (52.8) | 1950 (54.4)                 |
| Had cancer surgery since diagnosis                     | 16339 (45.1) | 15701 (43.3)                   | 1780 (49.6) | 1716 (47.9)                 |
| Mean hospital days (between 2 and 1 years before death)±SD | 0.82±1.12 | 0.84±1.17                     | 0.94±1.21 | 0.91±1.16                  |
| Disease duration                                        |        |                                 |         |                             |
| 0–5 years                                              | 28084 (77.5) | 29115 (80.3)                    | 2347 (65.4) | 2515 (70.1)                  |
| 6–11 years                                             | 4918 (13.6) | 4581 (12.6)                     | 656 (18.3) | 636 (17.7)                  |
| 12–17 years                                            | 2018 (5.6) | 1618 (4.5)                      | 324 (9.0)  | 249 (6.9)                   |
| 18+ years                                              | 1218 (3.4) | 924 (2.5)                       | 259 (7.2)  | 186 (5.2)                   |
| InterRAI Scales (When RAI-HC completed)                 |        |                                 |         |                             |
| Dependent on activities of daily living                | –      | –                               | 241 (6.7)  | 247 (6.9)                   |
| Minor-major depression                                 | –      | –                               | 496 (13.8) | 429 (10.0)                  |
| Moderate-to-severe cognitive impairment                | –      | –                               | 373 (10.4) | 363 (10.1)                  |
| Moderate-to-severe pain                                | –      | –                               | 391 (10.9) | 398 (11.1)                  |
| Caregiver present at Home                              | –      | –                               | 2279 (63.6) | 2264 (63.1)                 |

CHESS, Changes in Health, End-stage Disease, Signs and Symptoms; RAI-HC, Resident Assessment Instrument-Home Care.
DISCUSSION

In our population-based cohort of 114306 cancer decedents, a propensity score matched cohort of those who received palliative care earlier than 6 months before death compared with those who did not had a lower absolute risk of dying in hospital by 10%–13%, lower absolute risk of an aggressive care outcome in the last month of life by 10%, and higher absolute risk of having a supportive care outcome in the last month of life by 23%–38%. While prior randomised trials provided high internal validity within controlled settings, our approach provides high external validity in real-world settings. Bolstering the credibility that early palliative care is beneficial is the consistency of our findings across 2004–2014, which predate the publication of seminal randomised trials1–3; and the use of a population-based cancer cohort,

Table 2  Aggressive care measures in decedents with or without an RAI assessment

|               | NO-RAI |               | YES-RAI |               |
|---------------|--------|---------------|---------|---------------|
|               | Early palliative care | Not early palliative care | Absolute risk difference (%) (early vs not early)* | SD |
| Early palliative care | N=36238 (%) | N=36238 (%) | -10.0 | 0.21 |
| Not early palliative care | N=36238 (%) | N=36238 (%) | -10.4 | 0.21 |
| Absolute risk difference (%) (early vs not early)* | N=3586 (%) | N=3586 (%) | -13.2 | 0.21 |
| Death in acute care hospital | 13823 (38.1) | 17434 (48.1) | -10.0 | 0.21 |
| Aggressive care (any one or combination of the following three) | 18822 (51.9) | 22586 (62.3) | -10.4 | 0.21 |
| At least 1 ED visits within last 30 days | 15550 (42.9) | 19075 (52.6) | -9.7 | 0.21 |
| Any hospital admission within last 30 days | 16286 (44.9) | 19918 (55.0) | -10.1 | 0.25 |
| Any ICU admission within last 30 days | 1299 (3.6) | 2889 (8.0) | -4.4 | 0.85 |

*McNemar’s test was significant to <0.0001 for all measures.

ED, emergency department; RAI, Resident Assessment Instrument.

Table 3  Supportive home care measures in decedents with or without an RAI assessment

|               | NO-RAI |               | YES-RAI |               |
|---------------|--------|---------------|---------|---------------|
|               | Early palliative care | Not early palliative care | Absolute risk difference (%) (early vs not early)* | SD |
| Early palliative care | N=36238 (%) | N=36238 (%) | 23.3 | 0.39 |
| Not early palliative care | N=36238 (%) | N=36238 (%) | 22.3 | 0.73 |
| Absolute risk difference (%) (early vs not early)* | N=3586 (%) | N=3586 (%) | 37.8 | 0.39 |
| Supportive home care (any one or combination of the following three) | 22191 (61.2) | 13736 (37.9) | 23.3 | 0.39 |
| Physician house call in last 30 days | 9754 (26.9) | 6061 (16.7) | 10.2 | 0.86 |
| Palliative homecare nursing at home in last 30 days | 20370 (56.2) | 12320 (34.0) | 22.2 | 0.73 |
| Palliative personal support at home in last 30 days | 13728 (37.9) | 7954 (21.9) | 16.0 | 0.27 |

*McNemar’s test was significant to <0.0001 for all measures.

RAI, Resident Assessment Instrument.
meaning the findings were not a result of a particular cancer centre, intervention programme or cancer type.

This study addressed some of the noted gaps in prior research: it uses consistent exposure and outcome definitions over a long period of time and uses a large population-based cohort of all cancer types. Moreover, by using data from the RAI-HC, our study was able to control for previously unmeasured confounders known to be associated with receipt of early palliative care, such as worse pain, ADL dependency, depression, cognitive decline and health instability. This seeks to address selection bias in prior observational studies where those receiving palliative care might be different (eg, worse symptoms or have worse health instability) than those who do not. Our results were consistent with and without matching for RAI-HC variables. Moreover in our sensitivity analysis, where we examined early versus late subgroup separately (where the assumption was that both groups of patients were amendable to receiving palliative care), the findings were consistent with our overall study results, further supporting the benefits of early palliative care.

The results of this study support policies to enable earlier access to end-of-life homecare services and outpatient physician services for palliative care. In particular, policies that prohibit the access of palliative care services unless one forgoes curative treatments or is certified as expected to die within 6 months or less are disincentives to earlier and concurrent access to palliative care. For instance in the USA, the Medicare Hospice Benefit provides access to community-based hospice care but requires a physician to certify a life expectancy of less than 6 months and a patient commitment to forgo curative treatment. Besides policies, education is critical because research shows that patient preferences sometimes change over time, and that clinicians play an important role in introducing and initiating palliative care (eg, Serious Illness Conversations) and helping patients make informed treatment decisions about goals of care for end of life.

Our study has limitations. The propensity score matched design means we are comparing among those who are likely to have received early palliative care, but this may not represent the entire population of cancer decedents. We did not directly measure patient, family or provider preferences, which would be useful to control for in future studies. We used administrative data and billing codes to determine access to palliative care, which does not always represent the true intent of care provided, and we did not include billings from long-term care settings. As well, future research should examine outcomes of health system costs, health resource utilisation or patient and caregiver well-being.

In conclusion, across an 11-year population-based, cancer cohort, those who received early palliative care (before 6 months of death) compared with a matched cohort of those who did not, were more likely to receive supportive home care and less likely to receive hospital care in the last month of life. Our findings suggest that policies and education strategies to support the delivery of early palliative care might reduce the risk of dying in hospital and receiving aggressive care at end of life in real-world settings.
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