

Available online at www.sciencedirect.com**ScienceDirect**journal homepage: <http://www.elsevier.com/locate/rpor>**Original research article****Radiation therapy in the last month of life[☆]**

Anand Patel^a, Jacquelyn Dunmore-Griffith^a, Stephen Lutz^b,
Peter A.S. Johnstone^{c,*}

^a Department of Radiation Oncology, Howard University Hospital, Washington, DC 20001, United States

^b Department of Radiation Oncology, Blanchard Valley Health System, Findlay, OH 45840, United States

^c Department of Radiation Oncology, Indiana University School of Medicine, Indianapolis, IN 46202, United States

ARTICLE INFO**Article history:**

Received 19 June 2013

Received in revised form

23 August 2013

Accepted 23 September 2013

Keywords:

Health services research

Palliative care

Radiotherapy

ABSTRACT

Aim: We sought to survey a large, multi-center patient sample to better characterize/quantify RT utilization at the end of life.

Background: Few objective data exist for radiation therapy (RT) delivery at end of life (EOL).

Materials and methods: Data were retrieved for all patients receiving RT in calendar year 2010 in the Department of Radiation Oncology at Indiana University (IU) and Howard University (HU) hospitals. Specific attention was made of the group of patients receiving RT in the last 30 days of life.

Results: A total of 852 patients received all or part of their RT during 2010 (HU: 139, IU: 713). At time of analysis in early 2012, 179 patients had died (21%). Fifty-four patients (6.3% of total; 30% of expired patients) died within 30 days of receiving their last treatment. Twenty patients (2.3% of total; 11.2% of expired patients) received RT within their last week of life. For both sites, the median time until death from completion of therapy was 12.5 days (range 2–30 days).

Conclusions: Radiation in the last month of life is likely to provide minimal palliation or survival benefit. This, coupled with the financial implications, time investment, and physical costs, suggests that physicians and patients should more strongly consider hospice, and minimize duration of palliative RT courses as far as possible. As with chemotherapy, RT utilization at EOL should be considered for collection as an overuse metric.

© 2013 Greater Poland Cancer Centre. Published by Elsevier Urban & Partner Sp. z o.o. All rights reserved.

1. Background

Much of the modern practice of radiation therapy (RT) is palliative. Despite this, there are remarkably few data describing RT at end-of-life (EOL).^{1–5} With few national guidelines^{6,7} to unify medical opinion on such practice,

and minimal research describing efficacy of RT at EOL, it may be expected that there is a large variance in the treatment of terminally ill patients between different practices. In fact, one international survey suggested the existence of more than 100 different fractionation schemes used for patients treated with radiotherapy for bone metastases.⁸

[☆] Previously presented, in part, at American Radium Society Annual Conference, 5/2012, Las Vegas, NV, United States.

* Corresponding author at: Department of Radiation Oncology, Indiana University School of Medicine, 535 Barnhill Drive (RT041), Indianapolis, IN 46202, United States. Tel.: +1 317 944 2425; fax: +1 317 944 2486.

E-mail address: pajohnst@iupui.edu (P.A.S. Johnstone).

Hospice care aims to provide palliation of symptoms for patients whose expected disease course involves less than a 6 month life expectancy. Still, oncologists have sometimes been reluctant to refer patients for hospice consultation, and the average length of stay of oncology patients in hospice remains quite short.⁹ ASCO's Quality Oncology Practice Initiative (QOPI)¹⁰ has established a framework to guide management with chemotherapeutic agents in terminally ill patients. Chemotherapy provided within the last 2 weeks of life, as well as treatment courses initiated within 1 month of death are consistent with overutilization of chemotherapy. These practices were found to be associated with worse quality of life, with more intensive care unit admissions and more emergency department visits while failing to improve outcomes. One study compared a group of cancer patients including about 10% receiving chemotherapy at the EOL to another group of cancer patients containing 2% receiving EOL chemotherapy. The group with more EOL chemotherapy patients had a more than double rate of ICU admissions.^{11,12} Additionally, patients randomized to a palliative care consult at the time of diagnosis with non-small cell lung cancer tended to receive less chemotherapy and had a greater life expectancy than those who did not.¹³

No similar comprehensive scales like QOPI currently exist for RT. While chemotherapy at the end of life has been declared to be overaggressive, RT continued into or initiated during this timeframe has not been clearly discouraged in a similar fashion. Admittedly, a single fraction of 8 Gy may provide significant palliation of bone pain; but longer fractionation schemes likely have minimal additional palliative effect and confer no increase in survival. Thus, it is critical to analyze existing RT practices in terminal patients.

2. Aim

We undertook a benchmarking analysis to report the frequency of RT utilization for patients at EOL.

3. Materials and methods

To create a diverse sample set of patients encompassing a wide distribution of diagnoses, socioeconomic status, and stages of disease, all patients treated with RT in the 2010 calendar year were included in our cross-sectional study. The Institutional Review Boards at IU and HU provided approval for this retrospective analysis to be performed at those sites.

Several assumptions were made in interpreting these data. First, we assumed that radiation oncologists treated patients with the belief that they would benefit from RT. In other words, we assumed that no one was delivered RT considered a priori to be futile. Secondly, we assumed that briefer courses, generally of larger daily fraction sizes, were more frequently palliative than curative. This is of course subject to amendment based on the histology being treated. Frequency of palliative vs. definitive RT, sites of treatment, and specifics of concurrent chemotherapy, hormonal therapy or antibiotic therapy were specifically assumed to be similarly distributed between these two tertiary care facilities over the period of an entire year.

Table 1 – Summary of patient data.

	IU	HU
Total patients	713	139
Expired within 30 days of RT	44/6.2%	10/7.2%
Expired within 7 days of RT	16/2.2%	4/2.9%
Definitive cases dying within 30 days	4	3
Causes	MI (2); sepsis; prog dz	MI, sepsis, unk
Aborted courses	12/1.7%	5/3.6%
Mean fx delivered/prescribed	4.5/10	19/30

IU, Indiana University; HU, Howard University.

Prior to data collection, the sample's scope was narrowed to deceased patients only, and then among those who had deceased, only those who had passed away within 4 weeks of RT were considered for further analysis. Outcomes of interest that were defined prior to analysis were survival time post-treatment, oncologic diagnosis, and intention of treatment. Outcomes explored after collecting data included treatment timeframes, and received vs. prescribed dosage.

Dates of death for deceased patients were retrieved from the Social Security Death Index, and the date of death compared with the recorded last day of RT. Patients expiring within 30 days of receiving radiation therapy were selected for further analysis.

4. Results

Data are tabulated in Table 1. Briefly, at IU, 713 total patients received RT in 2010, of which 142 (19.9%) had died at time of analysis (summer, 2012). Of these 142 deceased, 44 patients (31%) died within 1 month of their last treatment. Of all 713 RT patients at IU in 2010, 6% died within a month of receiving treatment. At HU, 37 of the 139 total patients (26.6%) had died at time of analysis; ten of those 37 (27%) died within 1 month of their last treatment. Of all 139 RT patients at HU in 2010, 7% died within a month of receiving RT. There were no deaths due to RT toxicity.

At IU, of the 44 patients dying within 1 month of their last treatment, 16 patients (36.4%) received RT within their last week of life. At HU, of the ten patients who had expired within 30 days of RT, four patients (40%) received RT during the last week of life. The median time until death from completion of therapy was 12.5 days (range 2–30 days) at IU; at HU it was 12 days (range 3–29 days).

Definitive cases were less likely to die within 30 days of treatment. Four IU patients died while receiving definitive therapy: one patient undergoing chemoradiation for vulvar carcinoma died of sepsis, another died of small cell lung cancer, and two died of myocardial infarction. Three HU patients died while receiving definitive therapy.

At IU, 10 of the patients treated within 1 month of death had RT halted prematurely for worsening performance status, after a median 4.5 fractions of 10 prescribed. At HU, 4 EOL RT patients had RT halted prematurely for worsening performance status or for noncompliance, after a median

19 fractions of a median of 30 fractions prescribed (mean 27 Gy/45 Gy prescribed).

5. Conclusions

EOL issues are extraordinarily complex. Even something as conceptually simple as assigning “Do Not Resuscitate” status has vastly different ramifications for the patient, their loved ones, and health care team.¹⁴ The proportion of patients receiving chemotherapy in the United States within 14 days of death rose from 9.7% in 1993 to 11.6% in 1999.¹⁵ Justifiable concerns about this trend led to national initiatives such as QOPI to diminish what is perceived to be futile care. No such national data exist for RT. In determining appropriateness of RT at EOL, it is important to acknowledge the minimal palliative benefit and nonexistent survival benefit of radiating these patients. Only then, presuming a widening acceptance and use of hospice, can the national discussion begin about what aspects of RT at EOL may rightly be considered futile.

We previously published results of a pilot trial of patients presented at departmental Morbidity & Mortality (M&M) conference.¹ Of 63 patients receiving RT within 30 days of death, over half were still on treatment at time of death, and most of them were less than halfway through the prescribed course of RT. The inherent bias stemming from sampling from poor outcomes, wrought by this pilot study’s sample being selected from among M&M cases, compelled us to broaden the study population to include an entire year’s patients in a multi-institutional setting to provide a clearer perspective.

Gripp et al.² showed that among a set of patients receiving RT at the end of life, half of the patients actually had worsening symptoms despite palliative treatment intention. Only one quarter had improvement of their symptoms prior to death and the other quarter of the patients died while on treatment. In this group, half of cancer patients dying within 30 days of hospital admission had received daily radiation for more than 60% of their remaining life, and the median duration of RT was equal to the median survival of that cohort at 15 days. In our previous report on a select patient group,¹ over half (52%) of patients were still on treatment until the time of their death, with six patients receiving a treatment on the day of their death. Over half (54%) of patients had completed less than half of their original radiation therapy plan at the time of death and over one-fifth of patients were under treatment for more than half of their last month of life.

Kapadia and associates³ analyzed data between 2007 and 2010 from the National Comprehensive Cancer Network Non-Small Cell Lung Cancer Outcomes Database Project, reporting that 10% of patients who died had received EOL RT. Nearly half did not complete the prescribed course. Predictive factors for RT at EOL were stage IV disease or multiorgan involvement at diagnosis, age <65 years at diagnosis, and treating institution. The population of lung cancer patients was also the subject of a brief review by Ampil et al.⁴: specifically, they assessed whether the rapid decline after clinical detection of brain metastasis warrants immediate hospice referral rather than palliative RT.

Little else has been elucidated regarding what constitutes correct utilization of RT at EOL. Unlike chemotherapy, there

is no framework in place to validate halting radiation therapy either in the name of overutilization or futility. In order to decipher whether or not RT practices at the EOL are warranted, we first need to characterize the frequency of the practice, as we have done here. This study provides a look into the results among an unfiltered patient population receiving radiation therapy in the late stage of disease. Here we show that there is little inter-institutional variance in radiotherapy utilization at EOL between these two centers. In both patient groups the proportion of patients receiving radiation therapy within a month of death was 6–7%.

Conversely, some may argue that since only 6–7% of all patients treated in 2010 died within 30 days of RT, that 93–94% of patients did not, and that nothing needs change. The point of emphasis perhaps should be what cutoff is used: if we use RT within 7 days of death as a benchmark, 2–3% of patients treated in 2012 at these two sites meet that criterion. Determining what is ‘too many’ or ‘too few’ requires a national discussion and clearly is beyond the purview of this benchmarking manuscript. However, important issues that must be addressed include patients’ and their family’s acceptance of active treatment cessation in the context of palliative care, as well as physicians’ inexact ability to prognosticate the patient’s time of death.

It is unlikely that these data will quickly expand American hospice utilization to the last 6 months of life. As a start, we propose that physicians should consider ceasing therapy earlier in the terminal ill patient with worsening performance status and multiple signs of clinical deterioration, with the goal of transitioning care toward comfort measures and with the hope of avoiding ineffective treatment delivery in the last 4 weeks of life. From these data, we would propose to reduce that RT frequency from 6 to 7% of all patients receiving RT. The financial and physical costs involved mandate that physicians and patients should take greater advantage of hospice and work together to minimize duration of palliative courses as far as possible.

Conflict of interest

None declared.

Financial disclosure

None declared.

REFERENCES

1. Toole M, Lutz S, Johnstone PAS. Radiation oncology quality: aggressiveness of cancer care near the end-of-life. *J Am Coll Radiol* 2012;9:199–202.
2. Gripp S, Mjartan S, Boelke E, Willers R. Palliative radiotherapy tailored to life expectancy in end-stage cancer patients: reality or myth? *Cancer* 2010;116:3251–6.
3. Kapadia NS, Mamet R, Zornosa C, Niland JC, D’Amico TA, Hayman JA. Radiation therapy at the end of life in patients with incurable nonsmall cell lung cancer. *Cancer* 2012, <http://dx.doi.org/10.1002/cncr.27401> [Epub ahead of print].

4. Ampil F, Caldito G, Mills G, Marion J, Balandin A, Ponugupati J. Short survival after palliative radiotherapy for brain metastases in lung cancer: does the end justify the means? *Am J Hosp Palliat Care* 2012 [Epub ahead of print].
5. Lutz S, Korytko T, Nguyen J, et al. Palliative radiotherapy: when is it worth it and when is it not? *Cancer J* 2010;16: 473–82.
6. Lutz S, Berk L, Chang E, et al. Palliative radiotherapy for bone metastases: an ASTRO evidence based guideline. *Int J Radiat Oncol Biol Phys* 2011;79:965–76.
7. Tsao M, Rades D, Wirth A, et al. Radiotherapeutic and surgical management for newly diagnosed brain metastasis(es): an American Society for Radiation Oncology evidence-based guideline. *Pract Radiat Oncol* 2012;2: 210–25.
8. Fairchild A, Barnes E, Ghosh S, et al. International patterns of practice in palliative radiotherapy for painful bone metastases: evidence-based practice? *Int J Radiat Oncol Biol Phys* 2009;75:1501–10.
9. Von Gunten C, Lutz S, Ferris F. Why oncologists should refer earlier for hospice care. *Oncology* 2011;25:1278–80.
10. Neuss MN, Desch CE, McNiff KK, et al. A process for measuring the quality of cancer care: the quality oncology practice initiative. *J Clin Oncol* 2005;23:6233–9.
11. Ho TH, Barbera L, Saskin R, et al. Trends in the aggressiveness of end-of-life care in the universal health care system of Ontario, Canada. *J Clin Oncol* 2011;29:1587–91.
12. Barbera L, Paszat L, Chartier C, et al. Indicators of poor quality end-of-life cancer care in Ontario. *J Palliat Care* 2006;22:12–7.
13. Temel J, Greer J, Muzikansky A, et al. Early palliative care for patients with metastatic non-small-cell lung cancer. *N Engl J Med* 2010;363:733–42.
14. Trivedi S. Physician perspectives on resuscitation status and DNR order in elderly cancer patients. *Rep Pract Oncol Radiother* 2013;18:53–6.
15. Earle CC, Landrum MB, Souza JM, et al. Aggressiveness of cancer care near the end of life: is it a quality-of-care issue? *J Clin Oncol* 2008;26(23):3860–6.