Radiotherapy and Oncology xxx (2015) xxx-xxx



Contents lists available at ScienceDirect

Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com

Original article

The optimal utilization proportion of external beam radiotherapy in European countries: An ESTRO-HERO analysis $\stackrel{\circ}{\sim}$

Josep M. Borras^{a,*}, Yolande Lievens^b, Peter Dunscombe^c, Mary Coffey^d, Julian Malicki^e, Julieta Corral^{f,g}, Chiara Gasparotto^h, Noemie Defourny^h, Michael Bartonⁱ, Rob Verhoeven^j, Liesbeth van Eycken^k, Maja Primic-Zakelj¹, Maciej Trojanowski^m, Primoz Strojanⁿ, Cai Grau^o

^a University of Barcelona, Spain; ^b Ghent University Hospital, Belgium; ^c University of Calgary, Canada; ^d Trinity College, Dublin, Ireland; ^e Electroradiology Dep. University of Medical Sciences, Greater Poland Cancer Centre, Poznan, Poland; ^f Doctoral programme in Public Health, Department of Pediatrics, Obstetrics and Gynecology, Preventive Medicine and Public Health, Universitat Autònoma de Barcelona; ^g Catalonian Cancer Plan, Department of Health, Spain; ^h European Society for Radiotherapy and Oncology, Belgium; ⁱ CCORE, Ingham Institute for Applied Medical Research, UNSW, Liverpool, Australia; ^j The Netherlands Cancer Registry/Netherlands Comprehensive Cancer Organisation, The Netherlands; ^k Belgian Cancer Registry, Brussels, Belgium; ^l Cancer Registry, Institute of Oncology Ljubljana, Slovenia; ^o Carthus University Hospital, Denmark

ARTICLE INFO

Article history: Received 16 March 2015 Accepted 29 April 2015 Available online xxxx

Keywords: Radiotherapy Optimal utilization European countries Planning

ABSTRACT

Background and purpose: The absolute number of new cancer patients that will require at least one course of radiotherapy in each country of Europe was estimated.

Material and methods: The incidence and relative frequency of cancer types from the year 2012 European Cancer Observatory estimates were used in combination with the population-based stage at diagnosis from five cancer registries. These data were applied to the decision trees of the evidence-based indications to calculate the Optimal Utilization Proportion (OUP) by tumour site.

Results: In the minimum scenario, the OUP ranged from 47.0% in the Russian Federation to 53.2% in Belgium with no clear geographical pattern of the variability among countries. The impact of stage at diagnosis on the OUP by country was rather limited. Within the 24 countries where data on actual use of radiotherapy were available, a gap between optimal and actual use has been observed in most of the countries.

Conclusions: The actual utilization of radiotherapy is significantly lower than the optimal use predicted from the evidence based estimates in the literature. This discrepancy poses a major challenge for policy makers when planning the resources at the national level to improve the provision in European countries. © 2015 Elsevier Ireland Ltd. All rights reserved. Radiotherapy and Oncology xxx (2015) xxx-xxx

The estimated number of new cancer patients that require radiotherapy is a key parameter for planning the resources needed in a specific country in the framework of a cancer control programme. Most commonly, this calculation has been carried out using a specific proportion, typically the 'gold standard' of 50%, of the incident cases that would require radiotherapy at least

Radiotherap

00000 00000

* Corresponding author at: University of Barcelona – Hospital Duran i Reynals, Gran Via de l'Hospitalet, 199, 08908-Hospitalet del Llobregat, Barcelona, Spain. *E-mail address:* jmborras@iconcologia.net (J.M. Borras).

http://dx.doi.org/10.1016/j.radonc.2015.04.018 0167-8140/© 2015 Elsevier Ireland Ltd. All rights reserved.

^{*} Co-authors from HERO consortium: Arianit Kokobobo, University Hospital Center Mother Theresa, Tirana, Albania. Felix SedImayer, SALK and Paracelsus Medical University, Salzburg, Austria. Olivier De Hertogh, C.H. Peltzer – La Tourelle, Verviers, Belgium. Tatiana Hadjieva, University Hospital Queen Giovanna, Sofia, Bulgaria. Jiri Petera, University Hospital Hradec Kralove, Hradec Kralove, Czech Republic. Jesper Grau Eriksen, Odense University Hospital, Odense, Denmark. Jana Jaal, Tartu University Hospital, Tartu, Estonia. David Azria, CRLC Val d'Aurelle – Paul Lamarque, Montpellier, France. Zoltan Takacsi-Nagy, National Institute of Oncology, Budapest, Hungary. Jakob Johannsson, Landspitali University Hospital, Reykjavik, Iceland. Moya Cunningham, Luke's Radiation Oncology Network, St Luke's Hospital, Dublin, Ireland. Vydmantas Atkocius, National Cancer Institute, Vilnius, Lithuania. Michel Untereiner, Centre François Baclesse, Esch sur Alzette, Luxembourg. Martin Pirotta, Sir Paul Boffa Hospila, Floriana, Malta. Vanja Karadjinovic, Clinical Center Montenegro, Podgorica, Montenegro. Sverre Levernes, Norwegian Radiation Protection Authority, Osteras, Norway. Krystof Skladowski, Maria Sklodowska-Curie Memorial Cancer Center, Gliwice, Poland. Maria Lurdes Trigo, Instituto Português de Oncologia do Porto, Porto, Portugal. Barbara Šegedin, Institute of Oncology Ljubljana, Ljubljana, Slovenia. Sonia Garcia Cabezas, Hospital Universitario Reina Sofía, Cordoba, Spain. Magnus Lagerlund, University Hospital, Kalmar, Sweden. Bert Pastoors, Hôpitaux Universitaires de Genève, Geneva, Switzerland. Ben Slotman, VU University Medical Center, Amsterdam, The Netherlands. Charlotte Beardmore, Society and College of Radiographers, United Kingdom. Gaile Smyth, Belfast City Hospital, Belfast, Northern Ireland, United Kingdom. Jaap Vaarkamp, North Wales Cancer Treatment Centre (BCUHB), Bodelwyddan, Wales, United Kingdom. Agnieszka Dyzmann-Sroka, Cancer Registry, Greater Poland Cancer Centre, Poznan, Poland. Anna Kubiak, Cancer Regi

once during the course of his/her cancer. Although several more refined approaches have been developed to estimate this proportion developed by the Australian Collaboration for Cancer Outcomes Research and Evaluation (CCORE) can be considered the most optimal, as this approach involves a comprehensive and evidence-based analysis of all cancer sites [1] and has recently been updated [2]. In Australia this resulted in an optimal utilization proportion of 48.3%, that is, 48.3% of all incident cancer cases would require a course of radiotherapy at least once in the course of the disease.

In addition to the indications for radiotherapy in each clinical pathway, key parameters to evaluate the OUP of radiotherapy are the relative frequencies of cancer types as well as the stage at diagnosis [3].

The differences across European countries in cancer incidence, both in absolute number and in relative frequency of each tumour are significant [4]. Thus, it is of crucial importance to take into account the existing differences in the relative frequency of various cancers across Europe when calculating the OUP for each individual country, and to evaluate its impact on the absolute number of new cancer patients that will require radiotherapy.

The objective of this study is a logical step forward within the ESTRO-HERO project, which is an ESTRO supported activity aimed at developing a knowledge base and a model for the health economic evaluation of radiation oncology in the European countries. Analyses of the resources available in the European countries were recently published [5–7]. This paper evaluates the evidence-based country-specific demand for radiotherapy across Europe in order to allow all stakeholders to be able to estimate the capital and human resources required to deliver an appropriate radiotherapy service.

Material and methods

In order to estimate the absolute number of new cancer patients that will require radiotherapy in each country, the number of new cancer cases estimated for the year 2012, was combined with the overall OUP of radiotherapy in different European countries.

The country specific absolute number of new cancer cases has been obtained for the countries included in the European Cancer Observatory (ECO) database, estimated for the year 2012, and based on projections from the population based cancer registries available [4]. For non-Hodgkins lymphoma and head and neck cancers, which required further subdivision by subtypes, data from population based cancer registries have been used.

The OUP has been calculated using the methodology developed by the Australian Collaboration for Cancer Outcomes Research and Evaluation (CCORE) group, using evidence based indications for radiotherapy for all tumour sites with more than 1% incidence [1,2]. Briefly summarized, the CCORE team reviewed all the relevant published evidence-based guidelines and the scientific literature, updated until 2012, regarding the indications for radiotherapy for different tumour sites and accounting for the entire range of relevant stages at diagnosis. Indication for radiotherapy was defined as meaning it was the treatment of choice because there was evidence that radiotherapy has a superior clinical outcome (either measured by survival, quality of life, lower toxicity profile or better local control) compared to the alternative modalities or no treatment, provided that the patient is fit enough to undergo treatment. Based on these data, the CCORE team developed a decision tree model to estimate, by tumour site and for all cancers overall, the proportion of patients in whom external beam radiotherapy would be recommended at some stage during the course of the disease. Pathway probabilities included the distribution of cancer incidence by tumour site, stage at diagnosis and relevant clinical characteristics of patients for each tumour (age

and variations in performance status). The structure of the decision trees for each cancer site as well as the evidence supporting each clinical alternative and the corresponding probability of occurrence are available in the original report [8]. As neither retreatments nor brachytherapy were included in the CCORE decision trees, the focus of this analysis is exclusively on the optimal proportion of cancer patients receiving at least one course of external beam radiotherapy.

The relative frequency of each tumour is not only a necessary input for the calculation of the country-specific OUP, but also the main determinant of the inter-country OUP variation [3]. As shown in Table 1, which includes the relative percentages from the 12 most frequent cancers in Europe according to the ECO [4], the relative frequency varies significantly among countries. Stage at diagnosis, the other essential input parameter in the OUP calculation model, was not readily available in many cancer registries [9]. In a previous paper we have however shown that only a small part of the variability in overall OUP for radiotherapy was found among four European countries that have stage data available (i.e. Belgium, The Netherlands, Greater Poland region of Poland, and Slovenia), can be explained by the differences in stage distribution [3]. It can therefore be considered reasonable that the range of stage values from these four countries is representative of the other European countries. Hence, in order to assess the impact of stage distribution on the OUP in all ECO countries, we have applied the four different population based stage distributions, in addition to the Australian stage data, to each individual European country. The country specific range of OUP values are presented for all cancers combined

All OUP calculations are carried out using the year 2012 as reference for the incidence and frequency data. The available data for stage at diagnosis from the population cancer registries are for the years 2009–11, depending on the cancer registry [3].

The number of radiotherapy courses obtained by combining the estimated number of new cancer cases for 2012 with the OUP by country, can be considered the 'optimal demand for radiotherapy' in each individual country. This number has been compared to the annual courses of radiotherapy delivered from the HERO database (provided by the National Societies, available years ranging from 2009 to 2011; with the corrections of the data included) [5] in order to assess the gap between optimal and actual delivered treatment courses.

Data on the number of actual courses delivered were provided in different ways: without including retreatments in several countries while other countries included retreatments in the total number of courses. To allow comparison between optimal and actual courses, retreatments should be excluded. Therefore, in the countries where retreatments were included, the courses were adjusted by a factor of 0.80 in order to compensate for the increment of 25% typically applied for retreatments [5]. This approach has been taken due to the lack of consistent data across European countries on the retreatments carried out; until now only local analyses have been published [10-12], mostly from outside Europe. Thus, the option here has been to focus on data for new cancer cases. All calculations were carried out using the Tree Age software.

Results

The variability of overall OUP by country, adjusted by stage at diagnosis from the five population-based cancer registries, is presented in Fig. 1. The variation by country is typically limited to between 1% and 2%.

The numbers of new cancer cases estimated for the year 2012 in the 40 European countries included in this study are presented in Table 2, jointly with the calculated OUP by country. This OUP is

2

J.M. Borras et al. / Radiotherapy and Oncology xxx (2015) xxx-xxx

Table	1				
-			~		

Estimated percentage of	of cancer cases by	v most frequent tumour	site and country, 2012.
-------------------------	--------------------	------------------------	-------------------------

Country	Bladder	Breast	Corpus uteri	Kidney	Large bowel	Oral cavity	Lung	Melanoma of skin	Non-Hodgkin lymph	Pancreas	Prostate	Stomach	All sites
Albania	59	143	31	32	48	3.0	15.4	0.5	0.4	2.8	47	117	100.0
Austria	5.3	12.8	2.2	3.2	11.9	2.8	11.1	3.2	2.9	3.9	14.2	3.2	100.0
Belarus	3.3	11.7	4.4	4.9	12.3	3.8	12.4	1.8	1.6	2.5	6.6	9.1	100.0
Belgium	6.7	15.8	2.3	2.7	13.3	2.9	11.9	3.0	3.2	2.0	14.4	2.2	100.0
Bosnia	4.7	11.6	3.3	2.9	11.2	2.9	17.4	1.1	1.2	2.6	7.0	5.3	100.0
Bulgaria	5.2	12.3	4.0	2.7	15.4	2.7	12.3	1.4	1.7	3.9	5.7	5.2	100.0
Croatia	4.6	11.5	2.6	3.6	14.0	3.0	13.4	2.9	2.4	3.0	8.8	4.2	100.0
Cyprus	6.6	17.6	2.7	1.3	12.9	0.9	8.0	1.5	3.4	2.2	14.0	2.7	100.0
Czech Republic	4.3	11.9	3.3	5.7	14.5	2.6	11.6	3.8	2.2	3.7	11.9	2.8	100.0
Denmark	4.9	14.5	2.1	2.1	13.4	2.5	12.6	4.4	2.9	2.8	14.4	1.7	100.0
Estonia	3.4	10.8	3.4	4.6	12.9	2.3	10.3	2.7	2.1	3.1	16.7	6.0	100.0
Finland	3.8	15.7	3.0	3.1	10.2	2.0	8.8	4.2	4.2	4.0	18.9	2.3	100.0
France	3.0	14.6	1.8	3.0	11.0	3.0	10.8	2.7	3.1	2.5	19.8	1.8	100.0
FYR Macedonia	5.1	15.7	6.1	1.4	10.7	1.4	17.3	2.0	1.1	3.4	6.0	7.4	100.0
Germany	5.8	14.5	2.3	3.8	12.9	3.2	10.3	3.4	3.0	3.3	13.8	3.2	100.0
Greece	6.8	12.0	2.1	2.7	9.5	1.4	16.8	1.2	1.1	3.8	7.9	3.6	100.0
Hungary	5.3	10.1	1.6	3.1	16.7	5.3	18.4	2.2	2.0	3.7	6.3	3.9	100.0
Iceland	4.6	15.5	2.1	3.1	10.8	1.8	11.2	3.5	3.2	2.0	18.8	1.9	100.0
Ireland	3.2	13.9	1.8	2.7	12.3	1.8	10.9	4.1	3.4	2.5	18.2	2.3	100.0
Italy	5.2	14.3	2.4	3.2	13.6	1.6	10.5	2.8	3.5	3.0	12.6	3.7	100.0
Latvia	4.1	11.1	3.8	4.3	11.1	2.1	11.4	2.2	1.8	3.6	14.3	6.2	100.0
Lithuania	3.9	10.2	3.9	5.3	10.7	2.6	10.7	1.9	2.4	3.3	10.4	6.0	100.0
Luxembourg	3.9	14.5	4.6	2.8	12.5	2.6	10.5	3.5	2.7	2.7	13.6	2.7	100.0
Malta	6.9	16.5	3.4	3.0	14.1	2.7	9.5	1.9	2.6	3.9	10.6	3.6	100.0
Moldova	3.5	11.2	3.5	2.3	15.3	4.4	12.5	1.2	2.1	4.4	4.4	6.2	100.0
Montenegro	4.5	12.3	3.4	2.8	12.5	3.1	17.1	1.9	1.7	2.8	7.2	4.2	100.0
Norway	4.9	10.2	2.7	2.8	13.9	1.9	10.1	5.3	3.2	2.6	20.5	1.7	100.0
Poland	5.2	11.3	3.9	3.4	12.8	3.3	17.2	1.7	1.7	3.3	7.2	4.0	100.0
Portugal	5.8	12.4	3.0	2.0	14.5	4.2	8.5	2.2	3.7	2.5	13.5	6.1	100.0
Romania	4.9	11.4	2.0	2.5	13.0	4.7	14.8	1.4	2.0	3.9	5.8	5.2	100.0
Russia	3.0	12.5	4.6	4.2	13.1	3.2	12.2	1.9	1.7	3.2	5.9	8.4	100.0
Serbia	4.4	12.8	3.4	2.7	13.1	3.1	17.2	2.4	2.1	3.0	7.4	3.5	100.0
Slovakia	3.9	11.0	3.9	4.4	16.5	4.1	10.5	3.4	2.4	3.7	8.0	3.7	100.0
Slovenia	4.0	11.0	2.7	3.5	14.1	2.8	11.9	4.7	2.6	3.3	13.7	4.1	100.0
Spain	6.4	11.7	2.4	3.0	15.0	2.8	12.4	2.3	2.8	3.0	12.9	3.6	100.0
Sweden	4.7	13.1	2.8	2.2	12.6	1.9	7.7	5.8	3.2	1.9	23.0	1.6	100.0
Switzerland	5.0	13.7	2.4	2.3	11.6	2.8	10.1	5.9	3.6	2.8	18.7	1.6	100.0
The Netherlands	3.2	14.9	2.2	2.9	14.9	2.2	12.8	5.1	3.5	2.3	14.2	2.1	100.0
Ukraine	3.5	11.7	4.9	3.7	13.5	4.1	12.2	2.0	1.5	3.4	4.7	8.1	100.0
UK	2.7	16.0	2.6	3.0	12.4	2.3	12.3	4.4	3.6	2.7	13.9	2.0	100.0
Europe	4.4	13.5	2.9	3.3	13.0	2.9	11.9	2.9	2.7	3.0	12.1	4.1	100.0

Source: European Cancer Observatory (www.eco.iarc.fr/).

calculated by applying each stage data set, thus obtaining 5 different OUPs by country, and the highest and the lowest OUP are presented in the table.

Globally speaking, out of 3.41 million new cancer cases diagnosed in European countries in 2012, 1.74 million patients (unweighted average percentage of OUP between the highest and lowest stages of 51.0%) should have received at least one radiotherapy course following the evidence-based approach used in the present analysis. Using the stage distribution that provides the lower estimate of OUP, the variation in OUP by country ranged from the lowest in the Russian Federation with 47.0% to the highest in Belgium with 53.2%; or an absolute 6.2% difference is observed. No clear geographical pattern can be observed with respect to the OUP distribution.

The available numbers of radiotherapy courses by country from the HERO database are also presented in Table 2. The gap between the actual number and the optimal utilization obtained from the evidence-based model, expressed as a percentage, is calculated. This gap is presented in Fig. 2 as a percentage of the total number of patients that would have required, at least once, a radiotherapy course according to the OUP by country. Globally speaking, 4 countries treated at least 80% of the optimal indications for radiotherapy and 11 countries not even reached 70% of the patients optimally indicated. Of interest is that just one country reports a utilization of radiotherapy in excess to the figures proposed by the OUP range.

Discussion

The unique contribution of this study is the estimate of the optimal utilization proportion (OUP) of radiotherapy for 40 European countries. For each country, specific relative frequencies of the most common cancers were taken into account as well as realistic estimates of the ranges of stages at diagnosis from population based cancer registries. Previous attempts to calculate the number of new cancer cases that would require radiotherapy have only considered a unique proportion, usually 50% of new cancer cases plus 25% retreatments, 62.5% in total [13,14], all following the recommendations previously presented in the QUARTS project [15] that was based on the initial CCORE study.

Although the information on the distribution of cancer stages at diagnosis was not available for the majority of countries, which is a shortcoming, stage at diagnosis data from five population based cancer registries allowed us to take into account this second factor in the estimation of the OUP. The difficulties in collecting data on stage in population cancer registries are well known [9] and the number of registries with this information available is rather

Optimal utilization radiotherapy in Europe



Fig. 1. Range of values for overall optimal utilization proportion by country (in percentages of total cancer incidence).

limited. However, the four national sets of stage data together with those from Australia allowed us to create a range of values that could be considered as encompassing the majority of health services across Europe. In addition, we have assessed, in a previous paper, the differential impact of relative frequencies of tumours and stage distributions in five countries. The fact is that the relative frequency of tumours showed a higher impact on the OUP than stage distribution due to its more general influence on the decision trees [3]. The stage is very relevant to OUP in some specific tumour sites, such as rectal or cervical cancer, but not in others which significantly influence the demand for radiotherapy, such as breast or prostate cancer. Thus, it could be concluded that the lack of stage data for most countries has only a limited impact on the estimated national OUPs and the size of the impact can be judged from the rather small influence of the different stage distributions from the five countries for which data were available.

The approach applied here has shown that the variation in frequency distribution of individual cancer sites significantly affects the OUP calculation and hence the estimation of demand for radiotherapy with the concomitant resources - equipment and personnel - needed to cope with these new cancer cases. For instance, and using Belgium as an example, the OUP is 53.3%; using its own stage data [3]; a difference of 5.0% from the Australian estimate of 48.3% [2]. This difference represents 3267 additional patients with an indication for radiotherapy (excluding retreatments), implying the need for 7.8 linear accelerators, based on the average throughput of 420 courses of radiotherapy per MV unit in the European countries or 8.5 based on the Belgian average number of patients treated on a linear accelerator [5]. This illustrates how the differences calculated, even if they may seem small, could have a substantial impact on the calculation of the resources needed. The same example for Spain would result in an extra 7.2 linear accelerators. It can be concluded that the country-specific adjustment for the epidemiological situation by country may result in a more reliable estimate, which would provide a better basis to estimate the need for equipment and staff for radiotherapy, and consequently, more accurate planning at the national level.

The discrepancy between evidence based recommendations and the reality of the provision of health services poses a challenge for policy makers, although the range of discrepancies varies significantly among countries. The relevant question in this framework is why the actual utilization of radiotherapy is lower than it should be according to the evidence. In fact, several recent studies carried out in countries with high survival rates, a good indicator of quality of cancer care [16], such as the Netherlands or Norway, have shown that the proportion of patients receiving radiotherapy treatment is lower than expected. For instance, between 1997 and 2008, external beam radiotherapy was used in only 25% of the patients with a diagnosis of prostate cancer in the Netherlands [17], markedly lower than the OUP of about 60%, even taking into account that the utilization data from the Netherlands were restricted to the first 6 months after diagnosis. The situation was similar in non-small cell lung cancer [18] with 40% actual utilization, far below the calculated 80%; but it was not the case in rectal cancer, which showed a 71% of utilization [19] slightly higher than expected from CCORE model. Furthermore, in Norway [10], a lower utilization of radiotherapy during the first 5 years after diagnosis was found for all tumours in comparison with the evidence based recommendations. They also analyzed specifically the most frequent indications (breast, prostate, lung, colorectal, and head and neck cancers) and they observed a lower utilization than predicted, with the exception of breast cancer.

The discrepancy between evidence based OUPs and actual use of radiotherapy could be explained by different factors that are relevant when planning investments in radiotherapy resources.

- First of all, reduced use of radiotherapy has been associated with limitations due to geographical access [20], with specific impact on palliative radiotherapy [21].
- Second, the presence of comorbidity, particularly when combined with older age, is usually associated with the under-utilization of radiotherapy [22,23]. It should be recognized, however, that in specific indications such as muscle

4

J.M. Borras et al. / Radiotherapy and Oncology xxx (2015) xxx-xxx

Table 2

Summary of the assessment of incident cancer patients that will require radiotherapy treatment according to the evidence based utilization and excluding the need for retreatment.

Country	Total cancers $(n)^{a}$	Il cancers $(n)^a$ OUP $(\%)^b$		Optimal RT c	ourses (n)	Actual RT courses $(n)^{c}$	Actual/optimal RT courses (%)		
		Min.	Max.	OUP min.	OUP max.		OUP min.	OUP max.	
Albania	7143	52.6	54.3	3758	3879	2195	58.4	56.6	
Austria	41,117	49.0	50.3	20,155	20,698	17,185	85.3	83.0	
Belarus	32,422	48.5	50.3	15,738	16,293	NA			
Belgium	65,345	53.2	54.8 ^d	34,792	35,799	27,738	79.7	77.5	
Bosnia Herzegovina	9911	52.8	54.4	5236	5395	NA			
Bulgaria	32,053	51.3	53.0	16,434	16,977	11,035	67.1	65.0	
Croatia	22,890	51.2	52.7	11,717	12,055	NA			
Cyprus	3438	51.0	52.3	1753	1799	NA			
Czech Republic	57,627	48.5	50.2	27,943	28,945	26,104	93.4	90.2	
Denmark	36,119	52.8	54.3	19,064	19,600	14,144	74.2	72.2	
Estonia	6117	49.1	50.8	3004	3104	1698	56.5	54.7	
Finland	28,428	52.1	53.4	14,810	15,189	NA			
France	371,676	51.9	53.3	192,769	198,107	149,738	77.7	75.6	
FYR Macedonia	7330	52.6	54.3	3856	3981	NA			
Germany	493,780	50.1	51.6	247,419	254,735	NA			
Greece	40,971	52.5	54.2	21,523	22,213	NA			
Hungary	50,475	50.3	51.9	25,412	26,209	15,961	62.8	60.9	
Iceland	1449	50.7	51.8	734	750	476	64.8	63.5	
Ireland	20,808	51.5	52.9	10,714	11,017	6,698	62.5	60.8	
Italy	354,456	48.2	49.3	170,821	174,764	NA			
Latvia	10,347	49.9	51.4	5166	5315	NA			
Lithuania	14,520	49.9	51.5	7242	7483	5,014	69.2	67.0	
Luxembourg	2476	50.6	52.0	1252	1289	944	75.4	73.3	
Malta	1,902	51.9	53.3	988	1014	535	54.2	52.8	
Moldova	9894	50.2	52.1	4969	5151	NA			
Montenegro	2115	52.2	53.8	1105	1139	1200	108.6	105.4	
Netherlands	93,448	52.3	53.9 ^e	48,886	50,324	44,546	91.1	88.5	
Norway	28,214	49.0	50.5	13,818	14,248	10,786	78.1	75.7	
Poland	152,216	52.0	53.4 ^f	79,139	81,294	58,800	74.3	72.3	
Portugal	49,174	49.7	51.1	24,438	25,151	14,366	58.8	57.1	
Romania	78,760	50.0	51.8	39,383	40,805	NA			
Russian Federation	458,382	47.0	48.6	215,507	222,922	NA			
Serbia	42,221	52.2	53.8	22,050	22,733	NA			
Slovakia	24,045	48.2	50.2	11,599	12,071	NA			
Slovenia	11,457	49.6	51.3 ^g	5680	5874	3603	63.4	61.3	
Spain	215,534	49.7	51.1	107,018	110,159	78,820	73.7	71.6	
Sweden	50,481	51.4	52.8	25,928	26,662	18,142	70.0	68.0	
Switzerland	42,046	50.6	52.0	21,294	21,865	15,200	71.4	69.5	
Ukraine	140,999	50.2	52.1	70,811	73,403	NA			
United Kingdom	297,227 ^h	53.0	54.4	157,414	161,760	105,531 ⁱ	67.0	65.2	
Global	3,409,013	50.2	51.7	1,711,337	1,762,171				

NA: Not-available.

^a All cancers excl. non-melanoma skin cancer. Globocan 2012.

^b OUP: optimal utilization proportion.

^c Excluding re-treatment.

^d OUP calculated from population based stage at diagnosis from country cancer registry: 53.3%.

^e OUP calculated from population based stage at diagnosis from country cancer registry: 52.3%.

^f OUP calculated from population based stage at diagnosis from country cancer registry: 53.4%.

^g OUP calculated from population based stage at diagnosis from country cancer registry: 50.3%.

^h Scotland not included.

ⁱ Scotland not available.

invasive bladder cancer, where surgery is indicated, radiotherapy could be the preferred option with increased utilization in the elderly with comorbidity [24].

- Third, comparable effects were seen from patient-related factors such as the lower socio-economic level [25].
- A fourth relevant aspect is the preference of the physician for one therapeutic option over alternatives. In a paper published some years ago urologists and radiation oncologists were presented with the same clinical scenarios in order to assess their beliefs and therapeutic recommendations for prostate cancer. The result was not surprising that the specialists recommend the therapy that they deliver for the same clinical cases (72% of the radiation oncologists believed that external beam radiotherapy and surgery were equivalents while 93% of urologists believed that radical prostatectomy was the preferred option).

Beyond the specific clinical example, the paper clearly showed the bias favouring known clinical pathways that in some cases could exclude radiotherapy, although there is an evidence base for the indication [26]. Thus, lack of awareness, personal belief or knowledge could be another reason for the lower use of a particular treatment modality than expected considering evidence based data.

- A fifth factor is the shortage of resources, which resulted in waiting lists, delays in initiation of radiotherapy and reduced effectiveness of the treatment [20]. Such situations are unavoidably destinated to translate into lower utilization of radiotherapy. Shortage of resources is, at least in part, related to the specific financing structure within a country.
- This brings us to the last well-known factor that drives practice: the prevailing reimbursement in the country and the ensuing

Optimal utilization radiotherapy in Europe



Fig. 2. Comparison between actual and optimal utilization of radiotherapy by country (expressed as a percentage of the actual and the optimal number of patients, excluding retreatments).

financial (dis)incentives. In the case of radiotherapy, reimbursement mechanisms may play a role in the choice of treatment complexity or fractionation, as has been observed when the higher reimbursement of IMRT for prostate cancer fuelled its use in the US, or when fee-for-service reimbursement endorsed the use of more protracted schedules for the palliation of bone metastases, thus limiting the evidence-based use of single fractions [27,28]. Also, the fact that radiation oncology depends on the referral pattern of other specialists, adds complexity to this issue. But it is not impossible that, at the national level, the intricate interplay of the financing systems in oncology may result in radiotherapy being less attractive than competing oncologic treatment strategies.

Hence, translating evidence-based indications for radiotherapy into clinical practice requires taking into account all the above factors, with an emphasis on organizational factors related to accessibility and availability of capital resources and trained staff as well as the promotion of appropriate indications in a multidisciplinary framework of high quality cancer care, all endorsed by the appropriate financing mechanisms.

In order to fully assess these results, some limitations should be taken into account. The possible limitation of the OUP having been estimated with stage distributions from only five population based cancer registry data has been discussed above. Analyses, such as the one presented, are inherently limited by the timely and complete collection of data: for the estimates of the gap between actual and optimal utilization, the latter refers to the year 2012 while the data on actual utilization are not [5,6]. Variations in radiotherapy activity are, however, extremely dependent on facilities being opened or renovated and additional personnel being trained, both typically spreading over long time periods. Hence we feel that the impact of the differential timing of actual and calculated utilization on the estimated gap should not be too important, with possible local exceptions that could be included in the range of values estimated for each country. The last serious limitation is that the actual utilization is only available in about half of the countries. Thus, the discussion about the gap is limited to those countries

that have data on this key aspect of the planning of radiotherapy facilities and associated personnel. This point deserves careful consideration by the health care decision makers as the lack of essential data may seriously hamper rational planning of radiotherapy services.

In conclusion, the OUP for radiotherapy in the European countries has been estimated from an evidence-based assessment of the indications for radiotherapy, taking into account the differences in the relative frequency of cancer sites by country as well as a range of population based data on stage at diagnosis from five countries. A large discrepancy was observed between the actual utilization and the optimal utilization of radiotherapy in European countries, with less than 17% of countries treating at least 80% of the optimal indications for radiotherapy and about 46% of the European countries not even reaching 70% of the patients optimally indicated. These data should be taken into account when planning the resources at the national level and should support the development of guidelines for required resources and for cancer control plans.

Conflicts of interest

The authors have no conflict of interest.

Funding sources

This project was supported by the European Society for Radiotherapy and Oncology.

References

- Delaney G, Jacob S, Featherstone C, Barton M. The role of radiotherapy in cancer treatment: estimating optimal utilization from a review of evidencebased clinical guidelines. Cancer 2005;104:1129–37 (Erratum in: Cancer. 2006;107(3):660).
- [2] Barton MB, Jacob S, Shafiq J, et al. Estimating the demand for radiotherapy from the evidence: a review of changes from 2003 to 2012. Radiother Oncol 2014;112:140–4.

J.M. Borras et al./Radiotherapy and Oncology xxx (2015) xxx-xxx

- [3] Borràs JM, Barton MB, Grau C, et al. Impact of cancer incidence and stage on the optimal utilization of radiotherapy: methodology of a population based analysis by ESTRO-HERO project (2015, in press).
- [4] Ferlay J, Sterialova-Foucher E, Lortet-Tieulent J, et al. Cancer incidence and mortality patterns in Europe: estimates for 40 countries in 2012. Eur J Cancer 2013;49:1374–403.
- [5] Grau C, Defourney N, Malicki J, et al. Radiotherapy equipment and departments in the European countries: final results from the ESTRO-HERO survey. Radiother Oncol 2014;112:155–64.
- [6] Lievens Y, Defourny N, Coffey M, et al. Radiotherapy staffing in the European countries: final results from the ESTRO-HERO survey. Radiother Oncol 2014;112:178–86.
- [7] Dunscombe P, Grau C, Defourny N, et al. Guidelines for equipment and staffing of radiotherapy facilities in the European countries: final results of the ESTRO-HERO survey. Radiother Oncol 2014;112:165–77.
- [8] Ingham Institute for Applied Medical Research (IIAMR) Collaboration for Cancer Outcomes Research and Evaluation (CCORE). Review of optimal radiotherapy utilisation rates. CCORE report; 2013. Available from: tinyurl.com/pwkua34 [accessed 26-11-2014].
- [9] Siesling S, Kwast A, Gavin A, Baili P, Otter R. EUROCHIP-3 Workpackage 5. Availability of stage at diagnosis, cancer treatment delay and compliance with cancer guidelines as cancer registry indicators for cancer care in Europe: results of EUROCHIP-3 survey. Int J Cancer 2013;132:2910–7.
- [10] Asli L, Kvaloy S, Jetne V, et al. Utilization of radiation therapy in Norway after the implementation of the National cancer plan – A National, populationbased study. Int J Radiation Oncol Biol Phys 2014;90:707–14.
- [11] Khor R, Bressel M, Tai KH, et al. Patterns of retreatment with radiotherapy in a large academic centre. J Med Imaging Radiother Oncol 2013;57:610–6.
- [12] Barton MB, Allen S, Delaney GP, et al. Patterns of retreatment by radiotherapy. Clin Oncol 2014;26:611–8.
- [13] Rosenblatt E, Izewska J, Anacak Y, et al. Radiotherapy capacity in European countries: an analysis of the Directory of Radiotherapy Centres (DIRAC) database. Lancet Oncol 2013;14:e79–86.
- [14] Datta NR, Samei M, Bodis S. Radiotherapy infrastructure and human resources in Europe: present status and its implications for 2020. Eur J Cancer 2014;50:2735–43.
- [15] Bentzen S, Heeren G, Cottier B, et al. Towards evidence based guidelines for radiotherapy infrastructure and staffing needs in Europe: the ESTROQUARTS project. Radiother Oncol 2005;75:355–65.

- [16] De Angelis R, Sant M, Coleman MP, et al. Cancer survival in Europe 1999–2007 by country and age: results of EUROCARE–5-a population-based study. Lancet Oncol 2014;15:23–34.
- [17] Poortmans PMP, Aarts MJ, Jobsen JJ, et al. A population based study on the utilisation of primary radiotherapy for prostate cancer in 4 regions in the Netherlands, 1997–2008. Radiother Oncol 2011;99:207–13.
- [18] Koning CCE, Aarts MJ, Struikmans H, et al. Mapping the use of radiotherapy for patients with non-small cell lung cancer in the Netherlands between 1997– 2008. Clin Oncol 2012;24:e46–53.
- [19] Jobsen JJ, Aarts MJ, Siesling S, et al. Use of primary radiotherapy for rectal cancer in the Netherlands between 1997 and 2008: a population based study. Clin Oncol 2012;24:e1–8.
- [20] Mckillop WJ. Killing time: the consequences of delay in radiotherapy. Radiother Oncol 2007;84:1–4.
- [21] Popovic M, den Hartog M, Zhang L, et al. Review of international patterns of practice for the treatment of painful bone metastases with palliative radiotherapy from 1993 to 2013. Radiother Oncol 2014;111:11–7.
- [22] Vulto A, Louwman M, Rodrigus P, Coebergh JW. Referral rates and trends in radiotherapy as part of primary treatment of cancer in South Netherlands, 1998–2002. Radiolther Oncol 2006;78:131–7.
- [23] Chawla N, Butler EN, Lund J, Warren JL, Harlan LC, Yabroff KR. Patterns of colorectal cancer care in Europe, Australia and New Zealand. JNCI Monograph 2013;46:36–60.
- [24] Goossens-Laan C, Leliveld AM, Verhoeven R, et al. Effects of age and comorbidity on treatment and survival of patients with muscle invasive bladder cancer. Int J Cancer 2014;135:905–12.
- [25] Aarts MJ, Lemmens VE, Louwman MW, Kunst AE, Coebergh JW. Socioeconomic status and changing inequalities in colorectal cancer? A review of the association with risk, treatment and outcome. Eur J Cancer 2010;46: 2681–95.
- [26] Fowler F, Collins M, Albertsen P, et al. Comparison of recommendations by urologists and radiation oncologists for treatment of clinically localized prostate cancer. JAMA 2000;283:3217–22.
- [27] Jacobs BL, Zhang Y, Skolarus TA, Hollenbeck BK. Growth of high-cost intensitymodulated radiotherapy for prostate cancer raises concerns about overuse. Health Affairs (Project Hope) 2012;31:750–9.
- [28] Lievens Y, Van den Bogaert W, Rijnders A, Kutcher G, Kesteloot K. Palliative radiotherapy practice within Western European countries: impact of the radiotherapy financing system? Radiother Oncol 2000;56:289–95.