

Optimizing the Management of Prostate Cancer in Senior Adults: Call to Action

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INTRODUCTION

Prostate cancer is the most frequently diagnosed male cancer in developed countries [1]. With a total of 89,319 deaths in 2008, it represents the third leading cause of male cancer deaths in Europe, after lung and colorectal cancer [2]. It is the second leading cause of male cancer death in the U.S. [3]. Prostate cancer affects predominantly senior adults (i.e., men aged 75 years or older) [4]. According to the Surveillance Epidemiology and End Results Medicare database, median age at diagnosis in the U.S. is 67 years, and about two out of three deaths due to prostate cancer occur at the age of 75 years or older [4].

Life expectancy in the population as a whole is increasing [5], and we are seeing exponential aging worldwide. The number of people aged 80 years or older increased from 13.8 million in 1950 to 69.2 million in 2000, and a further increase to 379.0 million is forecast for 2050 [6]. Given this growth in the elderly population, the burden of prostate cancer is expected to worsen dramatically in the future. For example, it is anticipated that prostate cancer incidence in the U.S. will rise by 55% between 2010 and 2030, mainly driven by cancer diagnosis in older men, which is estimated to rise by 70% during the same period (Fig. 1) [7].

The management of senior adults with prostate cancer is not optimal. Only a minority of men over the age of 75 years with intermediate-risk or high-risk localized prostate cancer receive curative therapy (i.e., radical prostatectomy, radiation therapy, or brachytherapy), either in the U.S. (Fig. 2) [8] or in Europe [9].

The low rate of curative therapy in senior adults is attributed to the assumption that these older men will die from other



Figure 1. Projected new cases of prostate cancer per year in the U.S. (2010–2030) [7].

causes, not from prostate cancer. However, there is increasing evidence that men over the age of 70 years are more likely to have higher-risk prostate cancer—with higher clinical stage, Gleason score, and tumor volume—than younger men [10]. Without curative treatment, these aggressive tumors may lead to prostate cancer death in a significant number of men [11], which helps to explain the high rate of prostate cancer death at an advanced age. In a 20-year follow-up of a population-based cohort with localized prostate cancer between 1971 and 1984, men diagnosed after the age of 70 years of age with Gleason score 8–10 and managed conservatively (watchful waiting or androgen-deprivation therapy) had a 64% chance of dying of prostate cancer (Table 1) [12]. However, at an advanced stage of the disease, many oncologists are reluctant to give chemo-

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Figure 2. The age-specific percentage distribution of initial therapy by risk group [8]. Abbreviations: EBRT, external-beam radiotherapy; brachy, brachytherapy; RP, radical prostatectomy.

Table 1. Mortality due to prostate cancer or other causes in men aged 70–74 years with Gleason score 8–10 at dia	.gnosis
and managed conservatively [12]	-

20-year follow-up	Gleason score				
	2–4	5	6	7	8–10
Mortality due to prostate cancer (%)	12	14	28	41	64
Mortality due to other cause (%)	76	81	70	56	36
Missing data (%)	8	2	2	2	0
Missing data (%)	8	2	2	2	

therapy to prolong survival and improve patient's quality of life because of concerns over the tolerability.

The challenge for the physician is to accurately evaluate the life expectancy of a given patient and balance the risk of dying due to prostate cancer (which depends on the Gleason score and tumor stage) with the risk of dying due to other causes. Life-expectancy estimates provided by Social Security Administration (SSA) tables apply only to populations, not to individuals. Life expectancy is highly variable from one individual to another, and this variability reflects differences in health status. For example, median life expectancy of a 70-year old man in the SSA tables is 12.4 years, but healthy individuals are expected to live for at least 18 years (upper quartile), whereas those with comorbid conditions will live for only 6.7 years (Fig. 3) [13].

Health status in senior adults mainly depends on the number and severity of comorbidities, and this relationship has been clearly established in localized prostate cancer [14]. The Charlson index, which rates exclusively potentially lethal comorbid conditions [15], has been identified as a much more powerful predictor of nonprostate cancer death than chronological age [16]. Complications of curative therapies such as radical prostatectomy [11, 17] and brachytherapy [18] are also more related to Charlson index than age. Nevertheless, both the functional and the nutritional status of the patient are also important parameters to consider in health status evaluation because the degree of dependence in daily living activities and malnutrition have been shown to significantly influence survival [19, 20].

The International Society of Geriatric Oncology (SIOG) created a multidisciplinary Prostate Cancer Working Group to perform a systematic review of existing literature and establish recommendations for the management of senior adults with prostate cancer. The consensus of this task force is that senior men with prostate cancer should be managed according to their individual health status, which is mainly driven by the severity of associated comorbid conditions, not according to chronological age [14, 21]. Existing international recommendations [22–25] as well as national guidelines [26] form the backbone for the treatment of localized and advanced prostate cancer. However, it is important to supplement these recommendations with knowledge about the individual patient to determine which treatment strategy is most appropriate for that individual.

Best supportive care measures are also of primary importance to reduce the side effects of therapies and to optimize treatment outcomes, especially in senior adults who may be more affected by treatment complications compared with younger patients [21]. Quality of life is a key driver of treatment success at an advanced age, and it can be significantly improved by early detection and management of treatment side effects [14]. Therefore, active monitoring for any complications should be encouraged.

We can never stress enough the importance of multidisciplinary teams when deciding which treatment options to offer an individual patient. By bringing together surgeons, urologists, radiologists, social workers, and nurses, all of the available expertise can be combined, and varied aspects of individual patient care can be discussed in an evidence-based manner.

This supplement addresses all of these issues. The use of curative therapies—who should receive them and which treatment—is discussed by Heather Payne and Marcus Graefen. Matti Aapro details the emerging treatment opportunities for

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Figure 3. Life expectancy and health status in older men [13]. Adapted from Walter LC, Covinsky KE. Cancer screening in elderly patients: A framework for individualized decision making. JAMA 2001;285:2750–2756, with permission.

the management of advanced disease, and Florian Scotté presents his expertise in best supportive care to optimize the efficacy of therapies for senior adults with prostate cancer. The final article provides an overview of the SIOG recommendations for optimal management of prostate cancer in senior adults, which we hope will be useful in your daily practice.

This call to action aims to increase awareness of the growing burden of prostate cancer in senior adults and how improvements can be made to the way we currently treat many of these individuals. It is also a call to every country to systematically integrate training dedicated to senior adults in urology and oncology fellowship programs. Lastly, it is of primary importance that existing prostate cancer guidelines systematically integrate specific recommendations for the management of senior adults to optimize quality of care in this growing segment of the population.

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REFERENCES

1. GLOBOCAN 2008. Fast Stats: Estimated Age-Standardised Incidence and Mortality Rates. Geneva: World Health Organization, 2008.

2. Ferlay J, Parkin DM, Steliarova-Foucher E. Estimates of cancer incidence and mortality in Europe. Eur J Cancer 2010;46:765–781.

3. American Cancer Society. Cancer Facts & Figures 2012. Atlanta, GA: American Cancer Society, 2012.

4. Surveillance Epidemiology and End Results. SEER Stat Fact Sheets: Prostate. Bethesda, MD: National Cancer Institute, 2012.

5. Kochanek KD, Xu J, Murphy SL et al. Deaths: preliminary data for 2009. Natl Vital Stat Rep 2011;59:1– 51.

6. United Nations Department of Economic and Social Affairs Population Division. World population ageing: 1950–2050. Available at http://www.un.org/. Accessed June 1, 2012.

7. Smith BD, Smith GL, Hurria A et al. Future of cancer incidence in the United States: Burdens upon an aging, changing nation. J Clin Oncol 2009;27:2758–2765.

8. Hamilton AS, Albertsen PC, Johnson TK et al. Trends in the treatment of localized prostate cancer using supplemented cancer registry data. BJU Int 2010;107: 576–584.

9. Houterman S, Jannsen-Heijnen MLG, Vereiji CD et al. Greater influence of age than co-morbidity on primary treatments and complications of prostate cancer patients: An in-depth population-based study. Prostate Cancer Prostatic Dis 2006;9:179–184.

10. Sun L, Caire AA, Robertson CN et al. Men older than 70 years have higher risk prostate cancer and poorer survival in the early and late prostate specific antigen eras. J Urol 2009;182:2242–2249.

11. Sanchez-Salas R, Prapotnich D, Rozer F et al. Laparoscopic radical prostatectomy is feasible and effective in 'fit' senior adults with localized prostate cancer. BJU Int 2010;106:1530–1536.

12. Albertsen PC, Hanley JA, Fine J. 20-year outcomes following conservative management of clinically localized prostate cancer. JAMA 2005;293:2095–2101.

13. Walter LC, Covinsky KE. Cancer screening in elderly patients: A framework for individualized decision making. JAMA 2001;285:2750–2756.

14. Droz JP, Balducci L, Bolla M et al. Management of prostate cancer in older men: Recommendations of a working group of the International Society of Geriatric Oncology. BJU Int 2010;106:462–469.

15. Charlson ME, Pompei P, Ales KL et al. A new method of classifying prognostic comorbidity in longitudinal studies: Development and validation. J Chronic Dis 1987;40:373–383.

16. Tewari A, Johnson CC, Divine G et al. Long-term survival probability in men with clinically localized prostate cancer: A case-control, propensity modelling study stratified by race, age, treatment and comorbidities. J Urol 2004;171:1513–1519.

17. Begg CB, Riedel ER, Bach PB et al. Variations in morbidity after radical prostatectomy. N Engl J Med 2002;346:1138–1144.

18. Chen AB, D'Amico AV, Neville BA et al. Patient and treatment factors associated with complications after

prostate brachytherapy. J Clin Oncol 2006;24:5298-5304.

19. Rockwood K, Stadnyk K, MacKnight C. A brief clinical instrument to classify frailty in elderly people. Lancet 1999;353:205–206.

20. Blanc-Bisson C, Fonck M, Rainfray M. Undernutrition in elderly patients with cancer: target for diagnosis and intervention. Crit Rev Oncol Hematol 2008;67: 243–254.

21. Droz JP, Balducci L, Bolla M et al. Background for the proposal of SIOG guidelines for the management of prostate cancer in senior adults. Crit Rev Oncol Hematol 2010;73:68–91.

22. National Comprehensive Cancer Network. NCCN Clinical Practice Guidelines in Oncology: Prostate Cancer Version 3.2012. Available at http://www.nccn.org/. Accessed June 1, 2012.

23. Heidenreich A, Bastian PJ, Bellmunt J et al. Guidelines on prostate cancer. Arnhem: European Association of Urology, 2012. Available at http://www.uroweb.org/. Accessed June 1, 2012.

24. Mottet N, Bellmunt J, Bolla M et al. EAU guidelines on prostate cancer. Part II: Treatment of advanced, relapsing, and castration-resistant prostate cancer. Eur Urol 2011;59:572–583.

25. Thompson I, Thrasher JB, Aus G et al. Guideline for the management of clinically localised prostate cancer: 2007 update. J Urol 2007;177:2106–2131.

26. National Institute for Health and Clinical Excellence. Prostate cancer: Diagnosis and treatment. London: National Institute for Health and Clinical Excellence, 2008.