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Platinum Priority – Brief Correspondence

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Overall Survival in Patients with Localized Prostate Cancer in the US Veterans Health Administration: Is PIVOT Generalizable?

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Abstract

A better understanding of overall survival among patients with clinically localized prostate cancer (PCa) in the US Veterans Health Administration (VHA) is critical to inform PCa treatment decisions, especially in light of data from the Prostate Intervention Versus Observation Trial (PIVOT). We sought to describe patterns of survival for all patients with clinically localized PCa treated by the VHA. We created an analytic cohort of 35 954 patients with clinically localized PCa diagnosed from 1995 to 2001, approximating the PIVOT inclusion criteria (age of diagnosis ≤ 75 yr and clinical stage T2 or lower). Mean patient age was 65.9 yr, and median follow-up was 161 mo. Overall, 22.5% of patients were treated with surgery, 16.6% were treated with radiotherapy, and 23.1% were treated with androgen deprivation. Median survival of the entire cohort was 14 yr (25th, 75th percentiles, range: 7.9–20 yr). Among patients who received treatment with curative intent, median survival was 17.9 yr following surgery and 12.9 yr following radiotherapy. One-third of patients died within 10 yr of diagnosis compared with nearly half of the participants in PIVOT. This finding sounds a note of caution when generalizing the mortality data from PIVOT to VHA patients and those in the community.

Patient summary: More than one-third of patients diagnosed with clinically localized prostate cancer treated through the US Veterans Health Administration from 1995 to 2001 died within 10 yr of their diagnosis. Caution should be used when generalizing the estimates of competing mortality data from PIVOT.

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The comparative effectiveness of treatment for clinically localized prostate cancer (PCa) is widely debated [1]. Half or more of identified cases of PCa are clinically insignificant, as they are unlikely to cause symptoms or death [2]. The Prostate Intervention Versus Observation Trial (PIVOT)

randomized 731 men with localized PCa to surgery versus observation. Enrollees were diagnosed between 1994 and 2001, were from US Veterans Health Administration (VHA) and National Cancer Institute sites, were aged ≤ 75 yr, had serum prostate-specific antigen (PSA) < 50 ng/ml, had

biopsy-proven stage T1 or T2 PCa, and were deemed medically fit for radical prostatectomy, with an estimated life expectancy >10 yr. Survival was not different for patients treated with prostatectomy versus observation across the entire cohort [3].

Current clinical guidelines for PCa recommend using life expectancy estimates to guide the choice between definitive therapy and surveillance because PCa patients often die of competing causes [4]. Despite the selection of men thought to have significant longevity, nearly half (48.4%) of PIVOT participants had died after 10 yr of follow-up [3]. It is possible that competing risks of mortality might differ in the total VHA population, but little is known about the extent of that difference in patients with localized PCa.

The VHA operates the largest national integrated health care system in the United States [5]. The VHA cares for a predominantly male population, and PCa is the most common cancer diagnosis [6]. Users of the VHA represent a population with greater physical, mental, and social challenges than otherwise similar patients [7–11]; however, many older and multimorbid veterans with PCa receive treatment with curative intent [12]. We created a cohort of men with clinically localized PCa who met criteria similar to PIVOT inclusion criteria to better understand patterns of treatment and survival of patients with localized PCa in the VHA. We hypothesized that the survival of this national cohort would be lower than that reported in PIVOT.

We restricted our cohort to patients with PCa who were diagnosed in the period from 1995 through 2001, who were aged ≤75 yr, who had clinical stage T1 or T2 disease, and

who were in the Veterans Affairs Central Cancer Registry (VACCR) [6]. We excluded patients with clinical nodal or metastatic disease. The VACCR does not include PSA data for the study period; however, the vast majority of patients treated by the VHA have PSA levels <20 ng/ml [13]. The primary treatment was defined using mutually exclusive categories: surgery, radiotherapy, androgen deprivation therapy, or conservative management. We calculated survival using the US Department of Veterans Affairs Vital Status File, which compiles data from the VHA, the Centers for Medicare and Medicaid, the Social Security Administration, and the National Cemetery Association [14]. We censored patient follow-up as of January 1, 2015. We constructed Kaplan-Meier product limit estimates to determine median survival rates and survival rates at selected time points. We used SAS v9.4 (SAS Institute, Cary, NC, USA) and JMP Pro v12 (SAS Institute) for our analyses.

The mean age of our cohort was 65.9 yr, with 18 268 clinical stage T1 and 17 686 clinical stage T2 patients. Conservative management was the most common primary treatment ($n = 13\ 601$, 37.8%), followed by androgen deprivation therapy ($n = 8314$, 23.1%), surgery ($n = 8073$, 22.5%), and radiotherapy ($n = 5966$, 16.6%). Among the 12 630 patients aged ≥70 yr, 5607 received conservative management, 3815 received androgen deprivation therapy, 938 had surgery, and 2270 had radiotherapy (Fig. 1).

Median survival of the cohort was 14 yr (25th and 75th percentiles, range: 7.9–20 yr) and was similar for patients with clinical stage T1 and T2 cancers. Survival stratified by treatment type and age at diagnosis is shown in Figure 2.

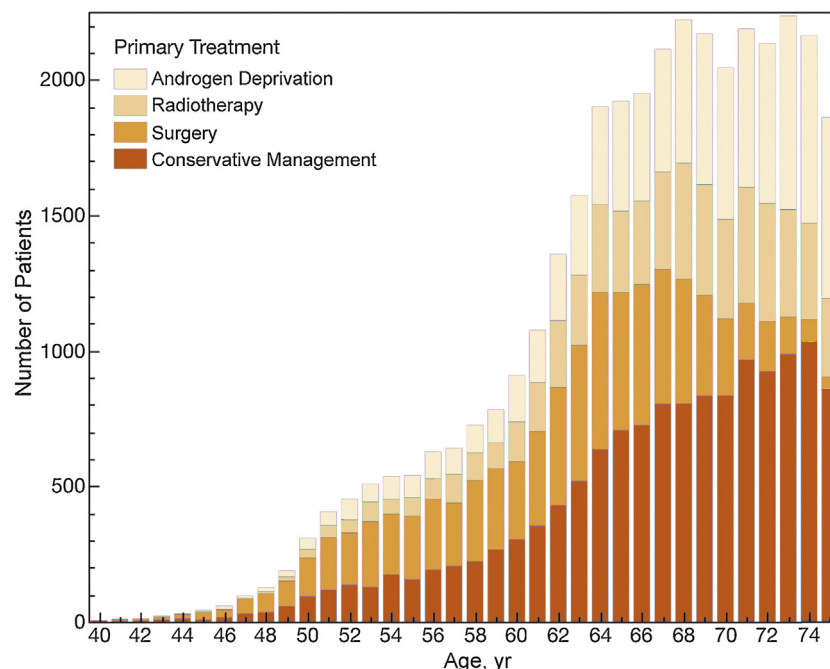


Fig. 1 – The primary treatment of clinically localized prostate cancer in the US Veterans Health Administration from 1995 to 2001. The total number of patients ($n = 35\ 954$) with clinically localized prostate cancer is plotted by patient age and treatment type. Surgery was the most common treatment among men aged <60 yr (2626, 42.8%) and declined steadily among men aged 60–69 yr (4509, 26.2%) and 70–75 yr (938, 7.4%). The percentage of men receiving radiotherapy increased slightly with increasing patient age (6.5% for age <50 yr, 12.7% for 50–59 yr, 17.2% for 60–69 yr, 18.0% for 70–75 yr). The number and percentage of patients treated with androgen deprivation therapy increased with patient age (11.7% for age <50 yr, 14.8% for 50–59 yr, 21.0% for 60–69 yr, 30.2% for 70–75 yr). Conservative therapy also increased with increasing patient age (30.3% for age <50 yr, 30.7% for 50–59 yr, 35.6% for 60–69 yr, 44.4% for 70–75 yr). Conservative management was the most common treatment for men aged >60 yr (39.3%).

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