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Technology diffusion and diagnostic testing for prostate cancer

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Abstract

Purpose—While the dissemination of robotic prostatectomy and intensity-modulated radiotherapy (IMRT) may fuel increased use of prostatectomy and radiotherapy, these new technologies may also have spillover effects related to diagnostic testing for prostate cancer. Therefore, we examined the association of regional technology penetration with receipt of prostate specific antigen (PSA) testing and prostate biopsy.

Methods—In this retrospective cohort study, we included 117,857 men age 66 and older from the 5% sample of Medicare beneficiaries living in the Surveillance Epidemiology and End Results (SEER) areas from 2003 – 2007. Regional technology penetration was measured as the number of providers performing robotic prostatectomy or IMRT per population in a healthcare market (i.e., hospital referral region). We assessed the association of technology penetration with rates of PSA testing and prostate biopsy with generalized estimating equations.

Results—High technology penetration was associated with increased rates of PSA testing (442 versus 425 per 1,000 person-years, p<0.01) and similar rates of prostate biopsy (10.1 versus 9.9 per 1,000 person-years, p=0.69). The impact of technology penetration on PSA testing and prostate biopsy was much smaller than the effect of age, race, and comorbidity (e.g., PSA testing rate per 1,000 person-years: 485 versus 373 for men with only one versus 3+ co-morbid conditions, p<0.01).

Conclusions—Increased technology penetration was associated with slightly higher rates of PSA testing and no change in prostate biopsy rates. Collectively, our findings temper concerns that adoption of new technology accelerates diagnostic testing for prostate cancer.

Keywords

intensity-modulated radiotherapy; prostate biopsy; prostate cancer; PSA testing; robotic prostatectomy

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Introduction

Prostate cancer is a common and expensive disease with annual spending exceeding \$11 billion in the United States in 2010.^{1,2} Treatment for prostate cancer is associated with significant morbidity, including urinary, sexual, and gastrointestinal side effects.³ With the promise of decreased morbidity and enhanced effectiveness, new therapeutic technologies such as intensity-modulated radiotherapy (IMRT) and robotic prostatectomy have been introduced over the last decade. However, both of these technologies carry significant price tags including start-up costs approaching \$3 million and, in the case of IMRT, high per episode costs.^{4–7}

Many worry that the imperative to recoup start-up costs for these expensive technologies fosters incentives to maximize their use.⁸ Acquisition of both IMRT and robotic capabilities was followed quickly by rapid increases in utilization of these treatment modalities, despite limited evidence supporting their effectiveness.^{7,9–11} While these direct effects of new therapeutic technologies are well established, there are no studies addressing their potential spillover effects on diagnostic practice patterns for prostate cancer. For instance, marketing and local press coverage of new technologies as well as specialist to primary care physician conversations may contribute to the perception of lower morbidity of treatment, which in turn may shift decision making in favor of identifying additional disease.^{12–14} Specifically, as the downsides of treatment appear lessened, local physicians may consciously or subconsciously alter their thresholds to recommend PSA testing or prostate biopsy.

For these reasons, we used linked SEER-Medicare data to evaluate the impact of marketlevel technology dissemination on diagnostic testing for prostate cancer.

Patients and Methods

Study population

To assess diagnostic testing for prostate cancer, we used the five percent Medicare sample to identify male beneficiaries residing in the Surveillance, Epidemiology, and End Results (SEER) areas between 2003 and 2007. This sample included men from the 5% non-cancer sample and men with cancer identified with the 5% flag. These men were followed beginning at age 66 years to allow assessment of their general health status in the year prior. Our study included only those without a prior diagnosis of prostate cancer enrolled in the Medicare fee-for-service program (i.e., eligible for Parts A and B of Medicare for at least 12 months and not enrolled in a Medicare Advantage plan, n=117,857). Subjects were followed for PSA testing and prostate biopsy until they were diagnosed with prostate cancer, lost Medicare eligibility, died, or until December 31, 2007, whichever came first.

To characterize prostate cancer technology diffusion, we used SEER – Medicare to identify all patients newly diagnosed with loco-regional prostate cancer between 2003 and 2007. We included subjects 66 years of age and older in the fee-for-service program eligible for Parts A and B of Medicare for at least 12 months before and after prostate cancer diagnosis (n=61,678).

Characterizing prostate cancer technology diffusion

We assessed technology diffusion for prostate cancer treatment at the level of a health care market, as defined by the Hospital Referral Region (HRR). Briefly, HRRs are a collection of ZIP codes in which Medicare beneficiaries residing in these areas receive their tertiary medical care.¹⁵ Medicare beneficiaries were assigned to their respective HRR based on the ZIP code of their primary residence.

We assessed regional prostate cancer technology penetration by measuring functional capacity, i.e., the per capita rates of physicians delivering robotic surgery or IMRT. Using explicit Healthcare Common Procedure Coding System (HCPCS) codes (see Table 1), we identified all patients who underwent robotic prostatectomy and IMRT.^{7,16} We then assigned each patient to a treating physician. The treating surgeon was identified using Unique Physician Identifier and National Provider Identifier Numbers, which are submitted with Medicare claims. The treating radiation oncologist was assigned in a similar manner as the provider who performed the clinical planning and simulation.¹⁷ We then assigned each physician to an HRR based on the provider ZIP code available in the physician claims.

Finally, we characterized regional technology penetration separately for robotic prostatectomy and IMRT for each year by calculating provider densities. The numerator was the number of physicians providing robotic prostatectomy or IMRT treatments in each HRR in a given year. The denominator was the number of male Medicare beneficiaries residing within the HRR based on population estimates for the ZIP Code Tabulation Areas.¹⁸ We then sorted HRRs into three equal groups (tertiles).

We validated our claims based measure of robotic prostatectomy providers by independently abstracting the number of providers offering this procedure in each HRR from the sole manufacturer's (Intuitive Surgical) historical webpages from 2004–2007. Historical websites were retrieved from web.archive.org using the search URL http:// www.davinciprostatectomy.com/hospitals.html on 09/27/2012. We found strong correlation between our claims based measure and the data abstracted from the historical webpages (r=0.81, p<0.001).

Outcome

Our primary outcome was the use of a diagnostic test to detect prostate cancer, either a PSA test or a prostate biopsy. For these measures, the numerators were whether or not at least one test was performed in each person-year; and the denominator was the time each subject was under observation in that year. We identified PSAs using explicit HCPCS codes in claims from the physician/carrier and outpatient files (see Table 1). To avoid counting PSA tests conducted in response to a suspicious test result, tests performed within 90 days of a prior test were excluded.¹⁹ Prostate biopsies were identified using claims from the physician/carrier file (Table 1).²⁰

Statistical analyses

Subjects were categorized by age, race, comorbidity,²¹ education, income, and urban residence (Table 2). Regional characteristics were obtained from the Health Resources and Services Administration's Area Resource File (number of hospital beds, number of urologists, and number of radiation oncologists per 100,000 men; Medicare managed care penetration) and categorized into tertiles based on the cohort. We described bivariate associations of demographic and regional characteristics with PSA testing and prostate biopsy by calculating the number of tests performed per person-year for each of the demographic or regional strata. To describe variation in PSA and prostate biopsy testing rates across HRRs, we calculated testing rates for each HRR and then described the median and range of these rates across all HRRs.

We performed bivariate and multivariable analyses with the person-year as our unit of analysis. The dependent variable was whether or not at least one PSA test was performed in a person-year. To account for the longitudinal nature of our data, we fit generalized estimating equations with a log link, using the natural logarithm of the time under observation as an offset.²² These models accounted for the fact that the likelihood of PSA

testing from one year to another is more similar within the same subject than between different subjects. They also allowed us to account for the nested structure of our data (i.e., subjects nested within HRRs). We then examined the association of regional technology penetration with PSA testing rates by adding the continuous measures of robotic prostatectomy and IMRT provider density to the models along with an interaction term. Technology penetration was allowed to vary from year to year within a given HRR. Multivariable models were adjusted for subject- and market-level covariates (neighborhood socioeconomic status,²³ number of hospital beds, number of urologists, and number of radiation oncologists per 100,000 men; Medicare managed care penetration, HRR-level provider volume), and for time in years since the beginning of the study. Because the effect of technology penetration on diagnostic testing could change over time, we also allowed for an interaction between technology penetration and time. From these models, we calculated adjusted PSA testing rates for the average HRR with low (i.e., robotic prostatectomy and IMRT penetration in the lowest tertile) and with high technology penetration (i.e., robotic prostatectomy and IMRT penetration in the highest tertile). The association of prostate biopsy rates with regional technology penetration was examined using similar models.

Sensitivity analyses

We performed sensitivity analyses excluding HRRs crossing SEER boundaries into areas from which SEER data was not available (n=21). We also estimated models not adjusting for number of urologists and radiation oncologists in a market, because adjusting for these covariates may be overly aggressive. We performed additional analyses only including PSA tests ordered by urologists or only including men less likely to benefit from prostate cancer diagnosis and treatment (i.e., those aged 75 and older with three or more comorbidities). The changes made in these sensitivity analyses did not materially affect the association between technology penetration and diagnostic testing for prostate cancer, so only results from the primary analyses are presented.

We performed all analyses using Stata version 12SE and SAS version 9.3. All tests were 2tailed; and we considered p<0.05 as statistically significant. The University of Michigan Medical School Institutional Review Board exempted this study from review in accordance with the Code of Federal Regulations Title 45, subpart A, section 46.101, paragraph b, subparagraph 4.

Results

Regional technology penetration for prostate cancer changed significantly over the study period for both robotic prostatectomy and IMRT (Figure 1, p<0.001). Typical low technology areas had a mean of 0 robotic prostatectomy and 5.7 IMRT providers compared with high technology areas that had a mean of 7.0 robotic prostatectomy and 17.6 IMRT providers per 100,000 male Medicare beneficiaries.

PSA testing and biopsy rates varied widely across HRRs, with a median of 391 (range 318 – 515) PSA tests per 1,000 person-years and 10.3 (range 3.7 – 19.3) prostate biopsies per 1,000 person-years. In bivariate analyses, older, more infirm, and more socioeconomically disadvantaged subjects had lower rates of PSA testing and prostate biopsy (Table 2). Regional technology penetration was associated with higher rates of PSA testing (Table 2). High IMRT penetration was associated with a small but statistically significant increase in prostate biopsy rates in bivariate analysis (Table 2). The effect of IMRT penetration was more pronounced in regions with high levels of surgical technology and the effect of robotic prostatectomy penetration was more pronounced in regions with high levels of IMRT technology (Table 3).

We accounted for this interaction between robotic prostatectomy and IMRT technology penetration in adjusted multivariable generalized estimating equations. Based on these models, living in an HRR with high technology penetration was associated with a small increase in the rates of PSA testing (from 425 to 442 per 1,000 person-years, p<0.001), while rates of prostate biopsy did not differ significantly (9.9 vs. 10.1 per 1,000 person-years, p=0.696, Figure 2). The impact of technology penetration on PSA testing and prostate biopsy was much smaller than the effect of age, race, and comorbidity (Figure 3).

Discussion

We found that PSA testing and prostate biopsy rates varied widely across health care markets (i.e., HRRs). Market-level technology penetration for prostate cancer treatment, measured as providers performing robotic prostatectomy and IMRT per capita, had the strongest effect on PSA testing in markets with both high robotic prostatectomy and high IMRT technology penetration. However, after adjusting for covariates in multivariable analyses, technology penetration had only minimal impact on PSA testing and no significant effect on prostate biopsy rates (Figure 2). The impact of technology penetration on diagnostic testing for prostate cancer was much smaller than the effect of immutable patient factors, such as age, race, and comorbidity (Figure 3).

This is the first study to broadly examine the impact of new technology on prostate cancerrelated services. Several studies found evidence that dissemination of robotic prostatectomy was associated with increased use of radical prostatectomy itself.^{9–11} For example, overall prostatectomy volume increased by a mean of 12 cases per year after a hospital acquired a robot while it decreased by 1 case per year for hospitals that did not acquire a robot between 2000 and 2009.¹¹ While these data provide evidence that dissemination of new technology impacts use of the technologies themselves, there could also be spillover effects related to diagnostic testing for prostate cancer. For example, pressure to recoup the high investment costs, marketing, and direct conversations among physicians^{12–14} may indirectly influence physician practices regarding PSA testing and prostate biopsies.

Therefore, we examined whether regional technology penetration for prostate cancer treatments had effects on diagnostic testing for prostate cancer. We found no effect on prostate biopsy rates and a statistically significant but clinically small effect on rates of PSA testing. For each one thousand patients followed for a year, only 17 additional PSA tests were performed in markets with high technology penetration. Our findings, to some degree, should allay concerns that these new technologies may spur the utilization of services aimed at identifying new cases of prostate cancer. Compared to technology penetration, patient factors such as age, race, and comorbidity were much more important predictors of PSA testing and prostate biopsy, which is consistent with previous studies examining factors associated with PSA testing.²⁴

Our study has several limitations. First, we could have incompletely ascertained the number of providers practicing in the 21 HRRs crossing SEER boundaries into areas from which SEER data was not available. Therefore, we performed sensitivity analyses excluding these fractional HRRs, with results that were consistent with our main findings. Second, given the limitations inherent to using Medicare data, our results may not reflect the effects of technology penetration on younger patients or those enrolled in Medicare Managed Care plans. However, the majority of prostate cancer diagnoses occur in the elderly²⁵ and men in this age group have the least to gain from early detection. Therefore, it is especially important to evaluate factors associated with PSA testing and prostate biopsy in this population. As such, our findings are generalizable to the largest incidence population at highest risk for overuse of PSA testing and prostate biopsy. Third, given the observational

nature of our data, we cannot exclude unmeasured confounding. To mitigate this problem, we controlled for a wide range of potential confounders, such as demographic and market factors, as well as socioeconomic class.

Conclusions

Our study has important implications for patients, payers, and policymakers. For patients, our findings provide insight regarding the degree to which the availability of new technology might influence the use of related healthcare services. For instance, when laparoscopic donor nephrectomy was introduced, there was a coincident significant increase in the number of individuals volunteering as living kidney donors,²⁶ likely because it increased the palatability of donation. We hypothesized that aggressive direct-to-consumer marketing of new technologies may increase the palatability of undergoing treatment for prostate cancer in a similar manner, which in turn could have upstream effects on PSA testing and prostate biopsy. However, our data suggest no clinically important association between dissemination of new technology and the use of these healthcare services, thus allaying concerns that dissemination of robotic prostatectomy and IMRT may drive diagnostic testing for prostate cancer. For payers and policymakers, our findings are of immediate interest as they consider coverage decisions for other new technologies such as proton beam therapy or histotripsy.^{27,28} While our work begins to elucidate the broader effects of technology dissemination on prostate cancer care, we will need future work evaluating the impact of new technology on the decision of whether or not to treat as well as on quality of care. This will allow us to gain a more complete understanding of how technology dissemination affects decision making along the prostate cancer care continuum.

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Abbreviations

HCPCS	Healthcare Common Procedure Coding System
HRR	Hospital Referral Region
IMRT	intensity-modulated radiotherapy

PSA	prostate specific antigen
SEER	Surveillance Epidemiology and End Results

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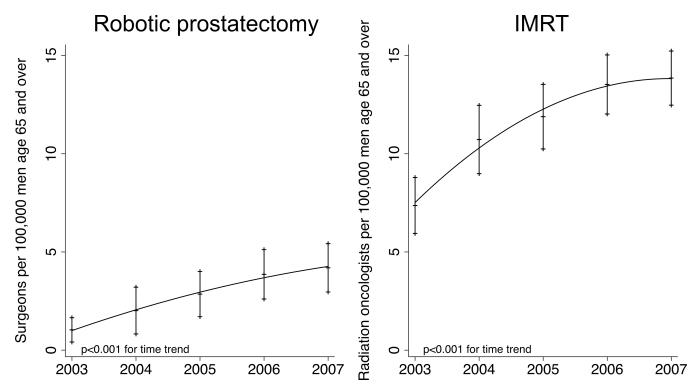


Figure 1.

Regional technology penetration for robotic prostatectomy and intensitymodulated radiotherapy (IMRT) across 69 Hospital Referral Regions in SEER-Medicare from 2003–2007.

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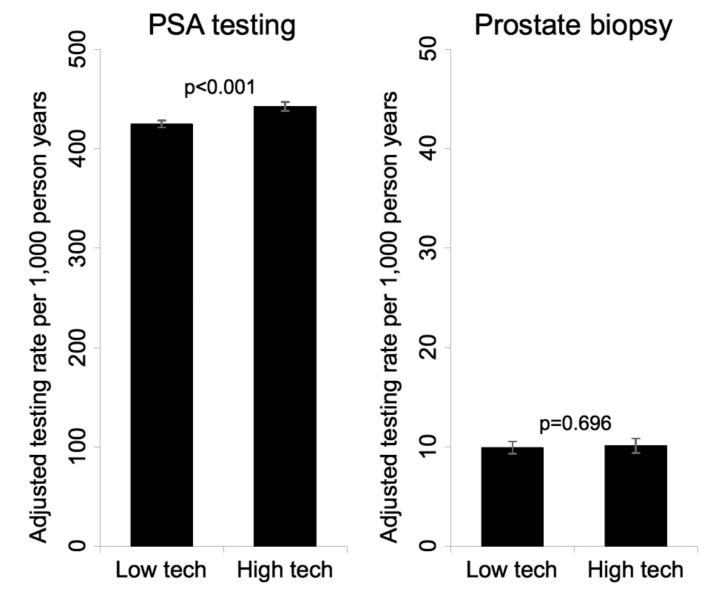


Figure 2.

Adjusted rates of PSA testing and prostate biopsy by regional technology penetration. Models were adjusted for socioeconomic status; rural vs. urban residence; number of hospital beds, number of urologists, and number of radiation oncologists per 100,000 men aged 65 and older; Medicare managed care penetration; market-level provider volume; and time in years since the beginning of the study. Error bars represent 95% confidence intervals.

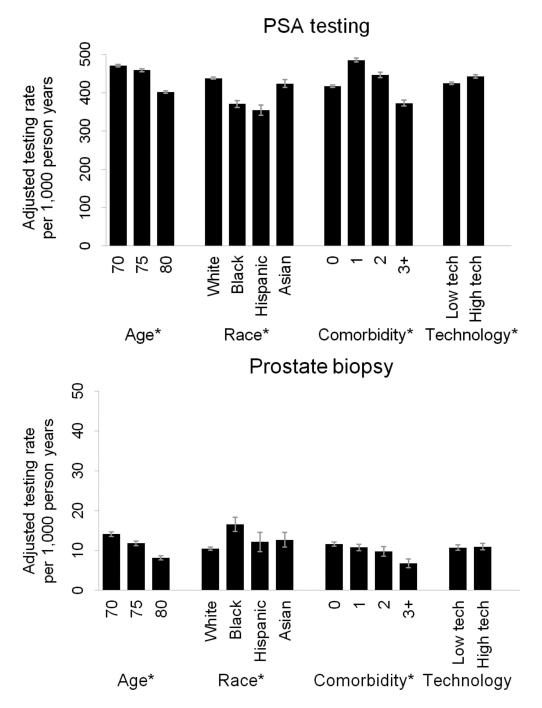


Figure 3.

Effect of regional technology penetration on PSA testing and prostate biopsy compared to effect of age, race, and comorbidity. Models were adjusted for socioeconomic status; rural vs. urban residence; number of hospital beds, number of urologists, and number of radiation oncologists per 100,000 men aged 65 and older; Medicare managed care penetration; market-level provider volume; and time in years since the beginning of the study. Error bars represent 95% confidence intervals. * denotes p<0.001.

Table 1

Healthcare Common Procedure Coding System (HCPCS) codes used in the analyses.

Procedure	HCPCS code		
Robotic prostatectomy*	55866		
IMRT	G0174, 77418, 0073T		
PSA testing	84153, G0103		
Prostate biopsy	55700		

*HCPCS code 55866 specifies laparoscopic radical prostatectomy, the overwhelming majority of which are known to be robotic prostatectomies.²⁹

Table 2

Rates of PSA testing or prostate biopsy. Numerator is the number of annual PSA tests or prostate biopsies, denominator is time in years subjects were under observation (subjects had to be alive, without prostate cancer, and eligible for both Part A and B at the beginning of a year to be included).

Characteristics	Person years	PSA testing rate per 1,000 person-years	Prostate biopsy rate per 1,000 person-years	
Age, years (%)*				
66–69	177,866	405	13	
70–74	104,056	462	13	
75–79	80,640	437	10	
80-84	50,922	369	7	
85+	29,041	261	3	
Race/ethnicity (%)*				
White	368,586	421	11	
Black	27,509	330	17	
Hispanic	12,023	314	11	
Asian	19,673	409	12	
Other/unknown	14,734	379	10	
Comorbidity (%) [*]				
0	291,681	403	12	
1	85,008	458	10	
2	36,569	418	9	
3+	29,267	343	6	
Lived in census tract in which 25% or more of adults had a college education **				
No	239,796	383	11	
Yes	188,901	448	12	
Median annual household income of census tract [*]				
Low (\$38,543)	141,960	366	10	
Intermediate	143,588	413	11	
High (\$54,091)	143,149	455	12	
Residing in rural area [*]				
No	374,758	419	11	
Yes	67,767	364	10	
Year ^{***}				
2003	89,247	407	12	
2004	89,319	399	11	
2005	88,277	406	10	
2006	87,641	413	11	
2007	88,041	428	11	

Characteristics	Person years	PSA testing rate per 1,000 person-years	Prostate biopsy rate per 1,000 person-years
Number of urologists per 100,000 men 65 and over st			
Low (55)	148,518	404	10
Intermediate	167,491	414	11
High (88)	126,516	414	12
Number of radiation oncologists per 100,000 men 65 and over st			
Low (23)	154,970	397	10
Intermediate	171,871	430	11
High (38)	115,684	400	12
Number of hospital beds per 100,000 men 65 and over ^{**}			
Low(4,797)	151,368	418	11
Intermediate	172,627	414	11
High (6,861)	118,530	397	12
Medicare managed care penetration **			
Low (5.2%)	149,293	400	11
Intermediate	152,538	420	12
High (21%)	140,694	411	10
Surgical technology****			
Low (0 providers per 100,000)	76,929	383	10
Intermediate	227,835	414	11
High (3 providers per 100,000)	137,761	420	11
IMRT technology**			
Low (8 providers per 100,000)	122,980	394	11
Intermediate	219,740	416	11
High (13.5 providers per 100,000)	99,805	418	12

IMRT=Intensity-modulated radiotherapy

P-values from bivariate generalized estimating equation models:

p < 0.001 for association with rates of PSA testing and biopsy

** p<0.001 for association with rates of PSA testing, p<0.05 for association with biopsy

*:

 $^{\ast\ast\ast}_{p<0.001}$ for association with rates of PSA testing, not significant for association with biopsy

Table 3

PSA testing rates (number of annual tests per 1,000 person-years) according to technology penetration.

IMRT technology penetration	Surgica	Total		
	Low	Intermediate	High	
Low	390	380	424	394
Intermediate	378	430	393	416
High	379	408	449	418
Total	383	414	420	411