

### ORIGINAL RESEARCH

# Early detection of prostate cancer using prostatespecific antigen testing: an empirical evaluation among general practitioners and urologists

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Background: Prostate cancer (PCa) is the most frequent cancer and the third leading cause of cancer death among German men. One option for PCa early detection is prostate-specific antigen (PSA) testing, which is still under debate regarding its risk benefits. Besides recommendations on the early PCa detection, daily practice on PSA testing varies in, for example, information communication and usage of the test. This pilot study assessed potential differences between general practitioners (GPs) and urologists in handling PSA testing and guidelines on early detection of PCa. Methods: 172 GPs belonging to the teaching network of the University of Oldenburg in Lower Saxony and Bremen and 128 practicing urologists were included in the online survey focusing on PSA testing. The questionnaire covered 43 questions on topics as the usage of the test, information communication, handling of test results and handling of/knowledge about national and international guidelines on PCa. Wether PSA testing is used in accordance with guidelines was also explored in four standardized case scenarios. Statistical analysis was done at a descriptive level. **Results:** In total, 65 doctors participated in the survey (response proportion: 21.7%, n=65; 27.9%, n=48 [GPs]; 13.2%, n=17 [urologists]). Results of 41 GPs and 14 urologists were analyzed. The PSA test was judged as useful by all urologists, while almost half of the GPs valued the test as ambivalent or not useful. Urologists showed a more proactive approach of informing men on PSA testing. Regarding guidelines and recommendations on PSA testing, GPs were less familiar with them compared to the urologists. Doctors of both specialties did not always treat men in consistence

Conclusion: This pilot study is highlighting differences in PSA testing practices between GPs and urologists in Germany. Urologists showed a more proactive approach. For further verification, we plan a more comprehensive study covering several German states.

Keywords: survey, guidelines, recommendations, testing practice, daily routine, PSA

with the guidelines. This was partially in contradiction to their self-appraisal.

## **Background**

Prostate cancer (PCa) incidence varies widely around the world, with by far the highest rates in North America and Oceania. In Germany, PCa is the most frequent cancer and the third leading cause of cancer death among men.<sup>2</sup> In the USA, the implementation of prostate-specific antigen (PSA) testing for the early detection of PCa in 1986 resulted in a rapid increase in PCa incidence. In Germany and other European countries, PSA testing was implemented later and increased in the 1990s.<sup>3</sup> Potentially caused by the introduction of the PSA testing, PCa incidence strongly increased during the following years in Germany, too. 4,5 Risks and benefits of this PCa screening method are controversial. Diagnosis at earlier stage is one potential advantage of the PSA testing,

Correspondence: Alexander Winter University Hospital for Urology, Klinikum Oldenburg, School of Medicine and Health Sciences, Carl von Ossietzky University Oldenburg, Rahel-Straus-Straße 10, Oldenburg 26133, Germany Tel +49 441 403 2302 Fax +49 441 403 2303 Email winter.alexander@klinikum-oldenburg. while disadvantages such as overdiagnosis and -treatment of harmless tumors followed by possible complications, eg, incontinence and impotence, also have to be considered.<sup>6,7</sup> Results of large studies such as the "European Randomized Study of Screening for Prostate Cancer" (ERSPC) and the "Prostate, Lung, Colorectal and Ovarian" (PLCO) trial show different findings. The ERSPC shows that the PCa mortality rates are possible to decrease by PSA testing, while in the PLCO trial they first concluded that screening was not associated with PCa mortality rates, whereas later results are in consistence with the ERSPC.8-11 Study results and recommendations based on these studies influence PSA testing practice and also are essential for policy making. 12-14 In contrast to breast cancer early detection, PSA testing has not been approved as a PCa early detection service of the German statutory health insurance. Therefore, in Germany, PSA testing is an individual health service which has to be paid by the patient himself.

The 2014 version of the German S3 guideline on PCa was the first one with a "dissenting opinion" by the German College of General Practitioners and Family Physicians (Deutsche Gesellschaft für Allgemeinmedizin und Familienmedizin [DEGAM]) for the use of PSA testing in the field of general practice. 15 Apart from differences in guidelines between countries and organizations, this guideline also states divergent recommendations for doctors working in different specialist disciplines, namely urology versus general practice. Urologists are advised to proactively inform men of at least 45 years of age and presumed life expectancy of at least 10 years of age about PSA testing, considering pros and cons. GPs should not actively inform men about PSA testing or only if the man asks for it. Therefore, it is expected that the daily routine in medical consultation, performance and other aspects concerning PSA testing varies. Despite the high significance of informed decision-making (for example emphasized in the German S3 guideline<sup>15</sup>), the clinical experience also shows that there is big variation in practice. The way the patient is informed has a high impact on the patient's demand and use of medical services as well as on the satisfaction with early detection by PSA testing. 16,17 A GP's recommendation clearly affects the patient's decision for or against a PSA test. 18 Furthermore, insufficient knowledge or individual conviction of the doctor influences treatment decisions in practice. 19,20 A survey among medical doctors with different specialties might highlight differences in PSA testing practice and in handling guidelines on PCa by specialist discipline.

This study was part of two affiliated pilot studies, aiming to assess attitude and practice of GPs and urologists on PSA testing. In this article, we focus on the survey based on the country-specific questionnaire covering the northwest of Lower Saxony and Bremen to investigate potential differences in daily routine practice regarding PSA testing between German GPs/internists and urologists.

### **Methods**

### Study population

A survey was set up to assess potential differences in routine PSA practice. The survey covered GPs within the teaching network of the University of Oldenburg in Lower Saxony and Bremen and urologists approached by the registry of the Professional Association of German Urologists (Berufsverband der Deutschen Urologen [BvDU]). We only considered practicing urologists for our study. Overall, 172 GPs and 128 urologists were invited to participate in the online survey.

### **Ouestionnaire**

The questionnaire covered 43 questions on several topics such as consultation, usage of the PSA test, processing of PSA test results, adherence to and knowledge about guidelines on PCa as well as characteristics of the respondents. Furthermore, the survey included four case scenarios to simulate situations and assess the GPs/urologist decision. The questionnaire was tested by three GPs and one urologist. The online questionnaire was developed using the software SoSci Survey (www.soscisurvey.de).

### Conducting the questionnaire

For conducting the questionnaire, the GPs and urologists first received an e-mail with information on the project and the upcoming questionnaire. A few weeks later (in June 2016), the survey started and the doctors received a link to the online questionnaire via e-mail. Three weeks later, the participants received a reminding e-mail. The interval for participation was about 7 weeks.

### Statistical analysis

Statistical analysis was done at descriptive level. We refer to the group urologists and the group GPs, where the latter also covers internists (n=5).

Results

### Response proportions and characteristics

Out of 172 GPs and 128 urologists, 64 participated overall, resulting in a response proportion of 21.3% (27.3%, n=47 [GPs]; 13.3%, n=17 [urologists]). Nine participants were excluded as they either did not finish the questionnaire (n=7) or were currently not working as a medical doctor (n=2). Unfortunately, no information is available for the nonparticipating physicians. Finally, the results of 55 questionnaires were considered for the analysis (see Figure 1). Group size differs between the two specialties (41 GPs and 14 urologists).

Tables 1 and 2 show the specialty and the characteristics of the respondents, respectively. The median age was 54.0 years for the GPs (IQR=10, mean=54.0, n=41) and 51.5 years for the urologists (IQR=8.75, mean=52.1, n=14). An unequal distribution was given for the sex of the respondents; there were no female urologists, while 87.8% (n=36) of the GPs were male (see Table 2).

### PSA testing practice

Questions on PSA testing are provided in Tables 3-6. More than two-third of the GPs (65.9%, n=27) and nearly 90.0% of the urologists (85.7%, n=12) stated to follow a standard procedure regarding PSA testing which is not older than 3 years for the majority (63.0%, n=17 [GPs]

and 58.3%, n=7 [urologists]). All urologists indicated that they inquire if the patient wishes to do a PSA test (majority of urologists [85.7%, n=12] orally). In the GP group, 24.4% (n=10) stated that they do not ask the patient if he wishes to do a PSA test, while 73.2% (n=30) do this orally (see Table 3).

Three-fourths of the GPs and all urologists always or often inform men on PSA testing during an early detection of cancer examination (75.6%, n=31 [GPs]; 100.0%, n=14 [urologists]). In case of discomfort in the lower urinary tract and unclear discomfort, GPs replies indicated a heterogeneous picture, whereas urologists showed a more proactive approach of informing men on PSA testing. Similar results can be found for the frequency of discussing certain factors with men before testing. The majority of all respondents stated to always or often discuss the listed factors (see Table 4). A more detailed table on the information communication of PSA testing can be found in Table S1.

One-third of the GPs and a proportion of 50% of the urologists answered on when a test is performed if a patient asks for it, that they either test the same day (36.6%, n=15 [GPs]; 57.1%, n=8 [urologists]) or cannot give a generalized answer ("depends on the patient") (34.1%, n=14 [GPs]; 42.9%, n=6 [urologists]).

More than half of the GPs (53.7%, n=22) replied that the proportion of men aged 45 years and older that finally receives (at least) one PSA test (irrespective of where the test is performed) is almost none or considerably less than

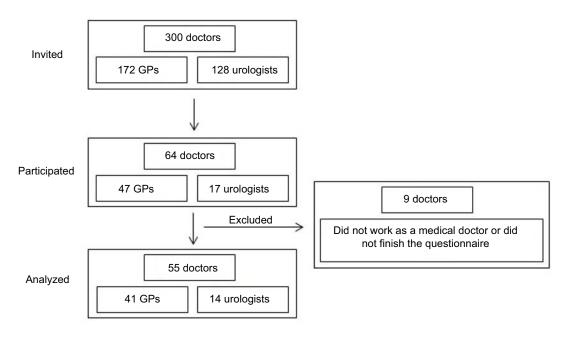


Figure I Flowchart of survey participants (general practitioners and urologists). Abbreviation: GPs, general practitioners.

Specialty	n	%
General medicine	31	56.4
General medicine and internal medicine	2	3.6
General medicine and surgery	2	3.6
General medicine and anaestesia	1	1.8
Internal medicine	5	9.1
Urology	14	25.5

**Table 2** Characteristics of respondents by general practitioners and urologists, n (%)

Variable	Categories	GPs	Urologists
Sex	Male	36 (87.8)	14 (100.0)
	Female	4 (9.8)	0 (0.0)
	Others	I (2.4)	0 (0.0)
Specialty	Yes	41 (100.0)	14 (100.0)
	No	0 (0.0)	0 (0.0)
Type of	Solo	12 (29.3)	4 (28.6)
practice	Group	23 (56.1)	8 (57.1)
	Practice sharing	5 (12.2)	1 (7.1)
	Community	I (2.4)	1 (7.1)
	health center		

Abbreviation: GPs, general practitioners.

half. On the other hand, almost 80% of the urologists (78.5%, n=11) stated that almost all or considerably more than half finally receive (at least) one PSA test, while none of them chose almost none or considerably less than half. Almost all GPs indicated that the blood sample is analyzed in an external laboratory (97.6%, n=40), while half of the urologists (50.0%, n=7) conduct the analysis in their own practice. Although almost 40.0% of the GPs (39.0%, n=15) would not recommend a test at all, the majority of the urologists (57.1%, n=8) chose 10–14 years of life expectancy for an asymptomatic patient to recommend a PSA test (see Table 5).

Figure 2 shows the boxplots of the number of PSA tests performed during the last 2 weeks by GPs and urologists. As expected, the median number of tests performed by urologists is higher than the corresponding one of the GPs (27.5, n=41 and 2.0, n=14, respectively).

Being asked which further actions were taken the last time seeing an asymptomatic patient with an increased PSA level, more than half of the GPs (53.7%, n=22) indicated to have checked the PSA level within a certain interval, while 68.3% (n=28) directly referred the patient to a urologist (see Table 6). All 14 urologists stated that

they checked the PSA level within a certain interval the last time they saw an asymptomatic patient with an increased PSA level.

Figure 3 shows the boxplots of the PSA level at which the participants would take further actions for an asymptomatic patient who received a PSA test within an early detection examination. For both groups the median was 4.0 (n=41 [GPs], n=14 [urologists]).

### Case scenarios

All urologists (100%, n=14) would recommend a PSA test to an asymptomatic patient without risk factors at a certain age. The corresponding figure for the GPs is 51.2% (n=21) (see Table 7, case scenario 1). Following up on case scenario 1, the minimum age is illustrated in Figure 4. All urologists (100%, n=14) stated to actively address a PSA test to a 45-year old patient with at least 10 years of life expectancy who does not ask for an early detection examination based on PSA testing. This statement was given by less than half of the GPs (41.5%, n=17) (see Table 7, case scenario 2). Almost one-third of the GPs (29.3%, n=12) would not recommend a second PSA test at all to a 45-year-old patient with a PSA level of 1-2 ng/mL (see Table 7, case scenario 3). More than one-third of the GPs (39.1%, n=16) would never or rarely perform a PSA test in a patient older than 45 years having an obstructive voiding disorder. All urologists would at least sometimes perform a test in such a situation, whereas the majority (50.0%, n=7) stated often (see Table 7, case scenario 4).

Figure 4 shows the boxplots of the minimum age at which GPs and urologists would recommend a PSA test to an asymptomatic patient of a certain age without risk factors. For both groups, the median was 45 years of age (n=41 [GPs], n=14 [urologists]).

### Guidelines and recommendations

Knowledge about national/international guidelines and studies on PSA testing was related to the specialist discipline of the doctors. Most GPs indicated to be aware of and consider the practice recommendations on PSA screening of the German College of General Practice and Family Medicine,<sup>21</sup> while all urologists indicated to know the German S3 guideline on PCa<sup>15</sup> in detail (see Table 8). Half of the GPs (48.8%, n=20) never heard about the European Association of Urology (EAU) guidelines on PCa.<sup>22</sup> International recommendations and guidelines

Table 3 Questions on PSA testing I by general practitioners and urologists, n (%)

Question	Categories	GPs	Urologists
Is there a standard procedure regarding PSA	Yes	27 (65.9)	12 (85.7)
testing (in your practice)?	No	14 (34.1)	2 (14.3)
If yes, how old is this standard?	$\leq$ 3 years	17 (63.0)	7 (58.3)
	4–9 years	6 (22.2)	5 (41.7)
	≥ 10 years	4 (14.8)	0 (0.0)
Who is responsible for medical consultation on	Doctor	11 (26.8)	2 (14.3)
PSA testing (in your practice)?	Doctor, medical assistant	3 (7.3)	0 (0.0)
	No reply	27 (65.9)	12 (85.7)
How do you ask the patient if there is a wish to	Not at all	10 (24.4)	0 (0.0)
do a PSA test?	Oral	30 (73.2)	12 (85.7)
	Standardized written form	0 (0.0)	1 (7.1)
	Oral, standardized written form	I (2.4)	1 (7.1)
How is the consultation on PSA testing done?	Oral	33 (80.5)	3 (21.4)
	Give away info material	0 (0.0)	1 (7.1)
	Oral, info material in waiting room	I (2.4)	5 (35.7)
	Oral, give away info material	7 (17.1)	3 (21.4)
	Oral, info material in waiting room and give away	0 (0.0)	2 (14.3)
	Others namely		
	- Online decision aid PSA (of the AOK)	I (2.4)	0 (0.0)
	- recommendations on info material regarding PSA	1 (2.4)	0 (0.0)
	testing		
	- internet links	I (2.4)	0 (0.0)

Abbreviations: AOK, Allgemeine Ortskrankenkasse, one of the German statutory health insurances; GPs, general practitioners; PSA, prostate-specific antigen.

such as the American Urological Association (AUA) guidelines on PCa,<sup>23</sup> the National Comprehensive Cancer Network (NCCN) guidelines on PCa early detection,<sup>24</sup> the American Cancer Society (ACS) guidelines on PCa early detection<sup>25</sup> and the recommendation statements of the US Preventive Services Task Force (USPSTF)<sup>26</sup> were rarely known in detail. Further, they were less known among GPs than among urologists. The same picture was given for international studies such as the ERSPC and PLCO study.<sup>8–11</sup> More than half of the GPs (58.5%, n=24) pointed out that they have never heard about the ERSPC.

Urologists mostly consider the German S3 and the EAU guideline(s) in their daily practice, while GPs indicated a rather moderate consideration of the DEGAM recommendations (see Table 9). None of the urologists and 5 (12.2%) of the GPs stated that their daily practice regarding PSA testing is less than moderately influenced by national/international studies and guidelines/recommendations (see Table 10).

Comparing the results of the questionnaire to the content of the guidelines, it can be noticed that urologists as well as GPs partly did not treat men in consistence with the guidelines. For example, >40% of the GPs (41.5%, n=17) would recommend a PSA test to an asymptomatic patient without risk factors, though both German guidelines stated for GPs not to recommend a PSA test to a patient described here (see Table 7, case scenario 2).<sup>21</sup>, On the question which interval doctors would recommend for a PSA test, seeing a 45-year-old patient with a life expectancy of at least 10 years, having a PSA level of 1–2ng/mL, less than half of the doctors chose the right answer according to the German S3 guideline (interval every 2 years; 36.6%, n=15 [GPs]; 42.9%, n=6 [urologists]) (see Table 7, case scenario 3).

## Daily practice

More than 80% of both groups always examine digito rectally during an early cancer detection examination (80.5%, n=33 [GPs] and 85.7%, n=12 [urologists]). A rather heterogeneous result was observed for patients with a voiding order – while all urologists (100%, n=14) always or often examine digito rectally, more than 40% of the GPs (41.5%, n=17) stated sometimes (see Table 11).

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 Table 4 Questions on PSA testing II by general practitioners and urologists, n (%)

Question	Categories	GPs			Urologists		
		Never + Rarely	Sometimes	Always + Often	Never + Rarely	Sometimes	Always + Often
On which occasions do you inform your patient on PSA	Within an early cancer detection	6 (14.6)	4 (9.8)	31 (75.6)	0.0) 0	0 (0.0)	14 (100.0)
testing?	examination						
	In case of a positive family anamnesis	6 (14.7)	4 (9.8)	31 (75.7)	0.0) 0	1 (7.1)	13 (92.9)
	In case of discomfort in the lower	16 (39.1)	8 (19.5)	17 (41.5)	I (7.1)	2 (14.3)	11 (78.5)
	urinary tract						
	In case of unclear discomfort	17 (41.5)	10 (24.4)	14 (34.2)	I (7.1)	3 (21.4)	10 (71.4)
How often do you discuss the following aspects with your	Impact on overall mortality	14 (34.2)	4 (9.8)	23 (56.1)	2 (14.2)	3 (21.4)	9 (64.3)
patients before performing a PSA test?	Impact on disease-specific mortality	7 (17.1)	3 (7.3)	31 (75.6)	2 (14.2)	1 (7.1)	11 (78.6)
	Impact on risk of metastasis	9 (22.0)	8 (19.5)	24 (58.6)	I (7.1)	1 (7.1)	12 (85.7)
	Potential overdiagnosis	3 (7.3)	5 (12.2)	33 (80.5)	2 (14.2)	1 (7.1)	11 (78.6)
	Issue of false positives	0.0) 0	6 (14.6)	35 (85.4)	0.0 (0)	0.0)	14 (100.0)
	Potential anxiety during waiting on	13 (31.7)	6 (14.6)	22 (53.7)	6 (42.9)	1 (7.1)	7 (50.0)
	test result						
	Potential follow-up examinations if the	1 (2.4)	8 (19.5)	32 (78.0)	0.0) 0	1 (7.1)	13 (92.8)
	test result is conspicuous						
	Adverse effects of the treatment	5 (12.2)	12 (29.3)	24 (58.5)	0.0) 0	6 (42.9)	8 (57.1)

Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen.

Table 5 Questions on PSA testing III, n (%)

Question	Categories	GPs	Urologists
When do you usually perform a PSA test if	Same day	15 (36.6)	8 (57.1)
a patient asks for it?	New appointment	9 (22.0)	0 (0.0)
	Depends on the patient	14 (34.1)	6 (42.9)
	Others, namely		0 (0.0)
	- after informing about benefit and risk	I (2.4)	-
	- test is only performed in justified exceptional cases	I (2.4)	-
	- sex, cycle	I (2.4)	-
Which proportion of men aged 45 years and	Almost none	7 (17.1)	0 (0.0)
older in your practice finally receives (at least)	Considerably less than half	15 (36.6)	0 (0.0)
one PSA test (irrespective of where the test is	Approximately half	8 (19.5)	3 (21.4)
performed)?	Considerably more than half	7 (17.1)	8 (57.1)
	Almost all	4 (9.8)	3 (21.4)
Where is the blood sample (of the PSA test)	In own practice	I (2.4)	7 (50.0)
analyzed?	External laboratory	40 (97.6)	6 (42.9)
	Others (eg, at a community health center)	0 (0.0)	1 (7.1)
How many years of life expectancy does an	Irrespective of the life expectancy (meaning also for	6 (14.6)	2 (14.3)
asymptomatic patient need to have at least for	patients with life expectancy of <5 years)		
you to recommend a PSA test?	5–9 years	5 (12.2)	4 (28.6)
	10-14 years	10 (24.4)	8 (57.1)
	≥ 15 years	4 (9.8)	0 (0.0)
	Not at all	16 (39.0)	0 (0.0)

Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen.

### PSA test and PCa risk

The general opinion on the PSA test differed between the groups (see Table 12). While two-third (64.3%, n=9) and one-third (35.7%, n=5) of the urologists judged the test as very useful and useful, respectively, the replies of the GPs were less homogeneous. More than one-third of the GPs (36.6%, n=15) judged the PSA test not to be useful. For a more detailed table on daily practice of digito rectal examining see Table S2.

Figure 5 shows the boxplots of the change in handling PSA testing during the last 10 years for GPs ( n=41) and urologists ( n=14). For the urologists, the median of 57.5 is slightly higher than the corresponding one of the GPs (53.0).

Figure 6 shows the boxplots of the score in agreement for undergoing a PSA test in the future for male GPs ( n=36) and male urologists ( n=14). The propensity to undergo a PSA test in the future is higher for urologists than for GPs (median 100.0 and 70.0, respectively).

The boxplots of the scores of GPs (n=41) and urologists (n=14) on the question if they think that the reduction of PCa-related mortality by early detection based on PSA testing is proven are shown in Figure 7. The median

score for the urologists is more than five times higher than the corresponding one for the GPs (19.0 and 100.5, respectively).

### Discussion

This study aimed to give a first insight into the daily practice of PSA testing and potential differences between GPs and urologists regarding the early detection of PCa based on PSA testing in the northwest of Germany. The results of this study show that GPs and urologists differ on various aspects regarding PSA testing. In total, PSA testing was less accepted among GPs than among urologists. Further, differences in opinion on usefulness of the PSA test, in handling PSA results, consideration of guidelines in daily practice and knowledge about guidelines or actual study results on PSA testing were observed.

Our findings support earlier studies showing that variation in handling of PSA testing is due to, for example, insufficient knowledge or individual conviction of the doctor. Also the fact that a doctors' intention to screen himself for PCa using PSA testing predicted their tendency to screen their patients is supported by our results. Because it is known that the way the patient is informed about the PSA test has a high impact on the

Table 6 Questions on PSA testing IV, n (%)

Question		Categories	GPs	Urologists
Which further actions did you	check the PSA level	Yes	22 (53.7)	14 (100)
take the last time having an	within a certain interval?			
asymptomatic patient with an		No	19 (46.3)	0 (0.0)
increased PSA level (according	directly refer the	Yes	28 (68.3)	n/a
to your definition of	patient to a urologist?			
increased)? Did you	(only GPs' replies)			
		No	13 (31.7)	n/a
	Others, namely	Sonography	I (2.4)	I (7.I)
		Rectal examination and sonography	I (2.4)	-
		Asked if the patient smokes	I (2.4)	-
		Consultation and wait-and-see attitude	I (2.4)	-
		PSA control after 6 months	-	I (7.I)
		PSA control after antibiosis	-	I (7.I)
Assuming you decided to	perform a third PSA	Yes	3 (21.3)	5 (35.7)
check the PSA level again	test?			
which, again, is conspicuous.		No	10 (76.9)	9 (64.3)
How did you proceed with your	directly refer the	Yes	8 (61.5)	n/a
last patient, again having an	patient to a urologist?			
increased PSA level? Did you	(only GPs' replies)			
		No	5 (38.5)	n/a
	Other, namely	Biopsy	-	5 (35.5)
		Biopsy or MRI of the prostate	-	I (7.I)
		Transrectal ultrasonography, if necessary	-	I (7.1)
		biopsy		
		Depending on the PSA level, if necessary	-	I (7.1)
		biopsy		
		Preclusion/treatment of infection	-	I (7.I)

Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen; n/a, not applicable.

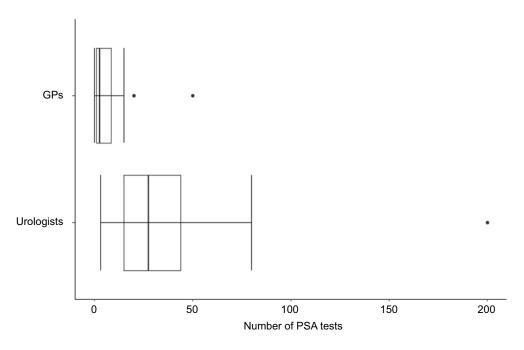


Figure 2 Number of PSA test performed during the last 2 weeks by general practitioners (n=41) and urologists (n=14). Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen.



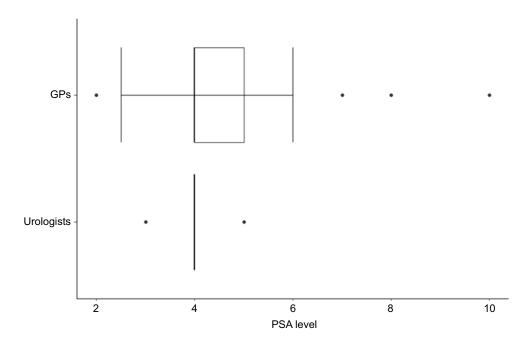


Figure 3 PSA level at which general practitioners (n=41) and urologists (n=14) would take further actions for an asymptomatic patient who received a PSA test within an early detection examination.

Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen.

Table 7 Case scenarios, n (%)

Question	Categories	GPs	Urologists
Case scenario I: Imagine you see an asymptomatic patient	Yes	21 (51.2)	14 (100.0)
without risk factors. Would you recommend him a PSA test at	No	19 (46.3)	0 (0.0)
a certain age?	Cannot reply to that question	I (2.4)	0 (0.0)
Case scenario 2: Imagine you see a 45-year old patient with life	Yes	17 (41.5)	14 (100.0)
expectancy of at least 10 years who does not ask for an early	No	24 (58.5)	0 (0.0)
detection examination based on PSA testing in your practice.			
Would you actively address a PSA test?			
Case scenario 3: Imagine a 45-year old patient with life expec-	Interval every year or more often	5 (12.2)	5 (35.7)
tancy of at least 10 years, having a PSA level of I-2ng/mL.	Interval every 2 years	15 (36.6)	6 (42.9)
Which interval would you recommend for a PSA test?	Interval every 3 years	I (2.4)	2 (14.3)
	Interval every 4 years	2 (4.9)	1 (7.1)
	Interval less than every 4 years	6 (14.6)	0 (0.0)
	Not at all	12 (29.3)	0 (0.0)
Case scenario 4: How often do you perform a PSA test in	Never	7 (17.1)	0 (0.0)
a patient older than 45 years having an obstructive voiding	Rarely	9 (22.0)	0 (0.0)
disorder?	Sometimes	13 (31.7)	4 (28.6)
	Often	8 (19.5)	7 (50.0)
	Always	4 (9.8)	3 (21.4)

Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen.

patient's satisfaction with early detection, 16 this indirectly influences their satisfaction with the urological care. One study conducted in the United States found no difference between urologists and primary care physicians in the amount of PSA tests conducted.<sup>29</sup> This is in

contrast to our findings. Differences in health care systems and/or opinions on PSA testing might be an explanation for this.

The results also show that doctors have a different self-assessment than the study results show. All



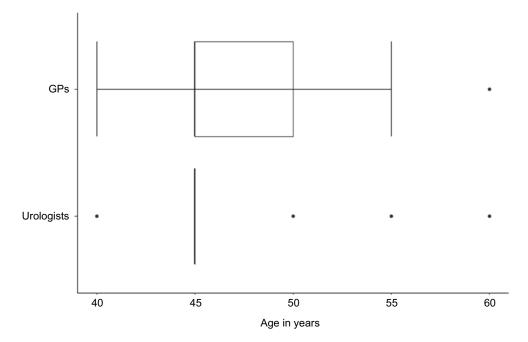


Figure 4 Case scenario 1a: Minimum age at which general practitioners (n=41) and urologists (n=14) would recommend a PSA test to an asymptomatic patient of a certain age without risk factors, if they would recommend one. Abbreviation: GPs, general practitioners.

Table 8 Questions on guidelines and recommendations I by general practitioners and urologists I, n (%)

Question	Categories	GPs			Urologists	;	
		Yes, I know it in detail	Yes, I heard about it	No, I never heard about it	Yes, I know it in detail	Yes, I heard about it	No, I never heard about it
Are you aware of the following guidelines	DEGAM	21 (51.2)	17 (41.5)	3 (7.3)	2 (14.3)	9 (64.3)	3 (21.4)
and study recommendations/results	recommenda-						
regarding PSA testing (irrespective of	tions						
the version)?	German S3 guideline	10 (24.4)	28 (68.3)	3 (7.3)	14 (100.0)	0 (0.0)	0 (0.0)
	EAU guidelines	3 (7.3)	18 (43.9)	20 (48.8)	10 (71.4)	4 (28.6)	0 (0.0)
	AUA recom- mendations	2 (4.9)	9 (22.0)	30 (73.2)	3 (21.4)	11 (78.6)	0 (0.0)
	NCCN guidelines	I (2.4)	7 (17.1)	33 (80.5)	I (7.I)	10 (71.4)	3 (21.4)
	ACS guidelines	I (2.4)	11 (26.8)	29 (70.7)	3 (21.4)	7 (50.0)	4 (28.6)
	USPSTF	I (2.4)	7 (17.1)	33 (80.5)	4 (28.6)	6 (42.9)	4 (28.6)
	recommenda- tions						
	ERSPC	2 (4.9)	15 (36.6)	24 (58.5)	7 (50.0)	5 (35.7)	2 (14.3)
	PLCO	2 (4.9)	7 (17.1)	32 (78.0)	4 (28.6)	5 (35.7)	5 (35.7)

Abbreviations: ACS, American Cancer Society; AUA, American Urological Association; EAU, European Association of Urology; DEGAM, German College of General Practice and Family Medicine; ERSPC, European Randomized Study of Screening for Prostate Cancer; GPs, general practitioners; PLCO, Prostate, Lung, Colorectal and Ovarian Cancer Screening Trial; PSA, prostate-specific antigen; NCCN, National Comprehensive Cancer Network; USPSTF, United States Preventive Services Task Force.

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Table 9 Questions on guidelines and recommendations II by general practitioners and urologists

Quartile 3rd 0.101 27.0 47.5 26.0 27.8 73.8 Quartile Urologists 9.18 26.5 12.8 5.0 <u>~</u> 0. <u>~</u> Median 20.5 34.5 73.5 19.0 12.5 6.0 5.0 26.8 (28.5) 13.1 (12.3) 21.3 (25.8) 13.4 (12.6) 42.8 (35.0) 10.8 (20.5) 11.8 (15.6) 64.8 (35.9) 94.9 (7.7) Mean (SD) 40 6 Quartile 3rd 83.8 46.0 20.0 9.8 18.5 29.0 37.0 Quartile st 6.0 <u>o.</u> <u>o</u>. 5.0 9. 9. Median 57.0 37.5 7.0 5.0 4.5 6.5 4.9 (19.1) 20.3 (22.5) 12.3 (16.0) 4.7 (17.4) 56.6 (31.4) 38.7 (28.9) 12.6 (17.7) 14.5 (17.7) 4.8 (18.8) Mean (SD) AUA recommendations German S3 guideline **Categories** recommendations recommendations **NCCNguidelines** EAU guidelines **ACS** guidelines DEGAM USPSTF ERSPC If yes (you are aware of the guidelines and study recommendations/results), do you consider them in your daily practice regarding PSA testing? Question

4 4 = 0 0

9 2

Abbreviations: ACS, American Cancer Society; AUA, American Urological Association; EAU, European Association of Urology; DEGAM, German College of General Practice and Family Medicine; ERSPC, European Randomized Study of Screening for Prostate Cancer; GPs, general practitioners; PLCO, Prostate, Lung, Colorectal and Ovarian Cancer Screening Trial; PSA, prostate-specific antigen; NCCN, National Comprehensive Cancer Network; USPSTF, United States Preventive Services Task Force.

**Table 10** Questions on guidelines and recommendations III by general practitioners and urologists, n (%)

Question	Categories	GPs	Urologists
To which extent do	Not at all	2	0 (0.0)
results of national and		(4.9)	
international studies	Very weak	2	0 (0.0)
and national and inter-		(4.9)	
national guidelines/	Weak	1	0 (0.0)
recommendations influ-		(2.4)	
ence your daily practice	Moderate	21	2 (12.5)
regarding PSA testing?		(51.2)	
	Strong	11	12 (85.7)
		(26.8)	
	Very strong	4	0 (0.0)
		(9.8)	

Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen.

urologists indicated to know the S3 guideline in detail and 94.9% stated to consider the guideline in their daily practice, while the answers of some questions were not in consistence with the guideline. This is in accordance with other studies stating that although doctors have favorable attitudes toward guidelines on PCa, guideline knowledge is limited.<sup>30,31</sup> One study stated that physicians handling after a normal or raised PSA seems to a large extent not in accordance with guidelines on PCa screening which agrees with our results.<sup>32</sup>

The "dissenting opinion" of the 2014 version of the German S3 guideline says that men, not broaching early PCa screening by PSA testing to the doctor, should not be actively approached by the GP, while urologists are advised to proactively inform these men about PSA testing. This might explain some differences between GPs and urologists, but not all. Apart from this "dissenting opinion" regarding a more reserved approach for GPs, the recommendations in the guideline (eg, retesting after a raised PSA, etc.) are identical for doctors with different specialties. Further, the results show that urologists know the urological guidelines better than GPs. However, this might not fully explain the observed differences between them.

Accordingly, there seems to be an urgent need to educate and support doctors more (GPs, as well as urologists and doctors with other specialties dealing with PSA testing) concerning PSA testing. <sup>19,27,30</sup> This is important for further improving the quality of the urological health care.

Table II Questions on daily practice by general practitioners and urologists, n (%)

Question	Categories	GPs			Urologis	sts	
		Never + Rarely	Sometimes	Always + Often	Never + Rarely	Sometimes	Always + Often
How often do you examine digito rectally in the following situations?	During an early cancer detection examination	0 (0.0)	4 (9.8)	37 (90.3)	0 (0.0)	0 (0.0)	14 (100.0)
	If there is blood in the patient's stool	0 (0.0)	3 (7.3)	38 (92.7)	(0.0)	0 (0.0)	14 (100.0)
	If the patient has a voiding disorder	4 (9.7)	17 (41.5)	20 (48.8)	0 (0.0)	0 (0.0)	14 (100.0)
	If the patient is asymptomatic	29 (70.3)	8 (19.5)	4 (9.8)	0 (0.0)	4 (28.6)	10 (71.5)

Abbreviation: GPs, general practitioners.

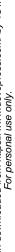
Table 12 Questions on PSA testing and prostate cancer risk, n (%)

Question	Categories	GPs	Urologists
How do you judge the PSA test in general?	Not useful at all	6 (14.6)	0 (0.0)
	Not useful	9 (22.0)	0 (0.0)
	Neither/nor	4 (9.8)	0 (0.0)
	Useful	18 (43.9)	5 (35.7)
	Very useful	4 (9.8)	9 (64.3)
Please state your opinion. Which of the following factors have an	Higher age (45 years and older)	41 (100.0)	14 (100.0)
impact on the risk to develop PCa?	Smoking	24 (58.5)	3 (21.4)
	Primary relative having PCa	32 (78.0)	13 (92.9)
	ВРН	5 (12.2)	2 (14.3)
	Afro-American ethnicity	2 (4.9)	9 (64.3)
Did you ever discover a PCa in an asymptomatic patient younger	Yes	29 (70.7)	14 (100.0)
than 60 years based on a PSA test that you performed?	No	12 (29.3)	0 (0.0)
Did you ever undergo a PSA test? (only men replies)	Yes	20 (55.6)	14 (100.0)
· · · · · · · · · · · · · · · · · · ·	No	16 (44.4)	0 (0.0)

Abbreviations: BPH, benign prostatic hyperplasia; GPs, general practitioners; PCa, prostate cancer, PSA, prostate-specific antigen.

In the future, non-PSA-based effective screening tests for early detection of PCa based on urine analysis may be more acceptable to the GPs because of probably improved specificity. Current studies aim the development of such innovative diagnostic tools which may improve the characterization of the disease biology, allowing to determine whether the PCa will be aggressive or indolent, in order to avoid overtreatment.<sup>33</sup> For example, urinary polyamines (eg, spermine) show potential to serve as novel PCa diagnostic biomarkers, which may be helpful to address the limited sensitivity and specificity problem of the serum PSA test.34

Certain limitations must be considered. One limitation is the fact that the GPs who were invited belonged to the teaching network of the University of Oldenburg. It is conceivable that the doctors belonging to this network are more interested in science. This selection bias could lead to an underestimation of the results. Although none of the urologists surveyed is currently working at a hospital, the urologists, in general, could have a different attitude toward PSA testing than GPs and other physicians due to their earlier training in clinical oncologic sites. The low response proportions of 27.9% for the GPs and 13.2% for the urologists are another limitation. However, other studies among doctors like GPs and



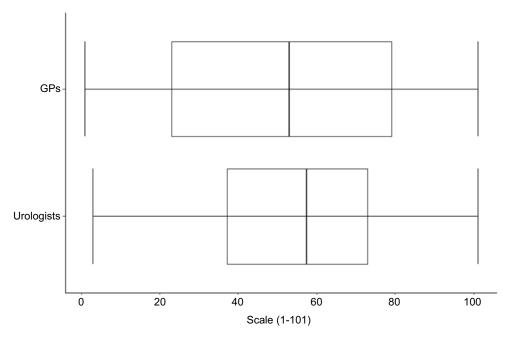


Figure 5 Change in handling PSA testing during the last 10 years for GPs (n=41) and urologists (n=14). Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen.

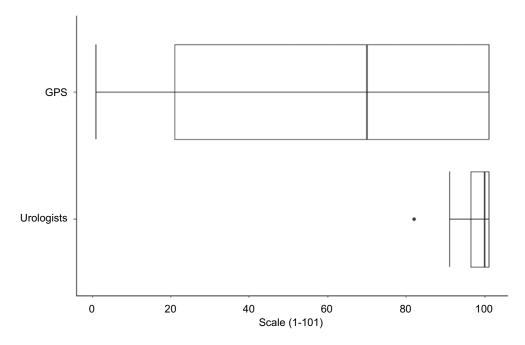


Figure 6 Score in agreement for undergoing a PSA test in the future for male general practitioners (GPs) (n=36) and male urologists (n=14). Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen.

psychiatrists show comparable response proportions. 35,36 A possible cause for this could be that physicians in Germany receive a lot of surveys, which partially are commercial and also well-paid. This survey was relatively extensive or time-consuming and without reimbursement. Thus, the participating physicians possibly had surpassing scientific interest and knowledge about clinical guidelines. As mentioned above, this circumstance may lead to selection bias.

Our pilot study provides an insight into the PSA tests conducted in the northwest of Germany. Till now, in Germany, only data from the German Health Interview

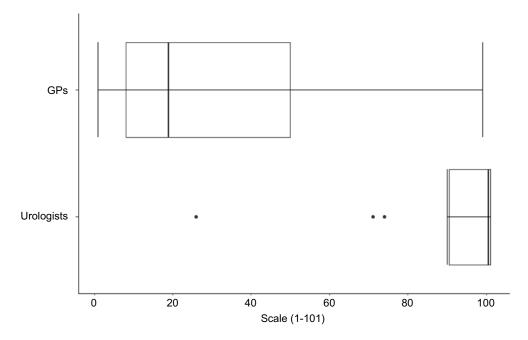


Figure 7 Scores of GPs (n=41) and urologists (n=14) on the question if they think that the reduction of PCa-related mortality by early detection based on PSA testing is proven. Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen...

and Examination Survey for Adults (2008-2011) are available, reporting that 30.6% of men aged 45 years or older had received a PSA test.<sup>37</sup> In other countries, these data often are determined based on secondary data.<sup>38</sup> As PSA testing is not an early detection service of the German statutory health insurance, these data are not gathered by the national health insurance.

If a patient wants a PSA test to be conducted, he has to pay the test by himself. The medical doctor "earns" a certain amount of money for every PSA test he or she conducts. Because financial incentives could affect doctors' treatment choices, this could be a motivation for a doctor to conduct a PSA test. 39,40 Our results also showed that 50.0% of the urologists analyzed the blood samples in their own practice. This could be an extra motivational factor for doctors to stimulate men to conduct a PSA test. To determine the influence of financial factors on conducting a PSA test further research is needed.

### **Conclusion**

This pilot study argues for differences in various aspects regarding PSA testing between GPs and urologists, wherein a low response rate represents a limitation in some respects. Physicians with surpassing scientific interest are probably over-represented, which may lead to selection bias. There is an urgent need to educate and support doctors more on topics related to PSA testing. This is important for further improving the quality of urological health care. Therefore, the interdisciplinary exchange needs to be continued and extended to achieve a consistent level of knowledge among doctors with different specialties.

To validate the results of this study and to constitute the consequences of the different information levels on the urological care, a follow-up project is planned. This project will be conducted in different regions in Germany and will include GPs as well as practicing urologists and those working at hospitals. Further, a survey among men aged >45 years is planned to assess the satisfaction with the urological health care in Germany.

## Ethics approval and consent to participate

The study was approved by the Medical Ethics Committee of the Carl von Ossietzky University Oldenburg (No. 041/2016).

### Availability of data and materials

Data and materials supporting the conclusion were included in the main paper. Further data were available from the corresponding author on reasonable request.

### **Abbreviation list**

ACS, American Cancer Society; ASR, Age-standardized rate; AUA, American Urological Association; BvDU, Professional Association German **Urologists** of (Berufsverband der Deutschen Urologen); BPH, benign prostatic hyperplasia; DEGAM, German College of General Practice and Family Medicine (Deutsche Gesellschaft für Allgemeinmedizin und Familienmedizin); EAU, European Association of Urology; ERSPC, European Randomized Study Screening for Prostate Cancer; GP, General practitioner; NCCN, National Comprehensive Cancer Network; ng/mL, nanograms per milliliter; PCa, prostate cancer; PLCO, Prostate, Lung, Colorectal and Ovarian study; PSA, prostate-specific antigen; USPSTF, US Preventive Services Task Force.

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### **Author contributions**

VJ and AW were responsible for the study design. SK, VJ and AW did the literature search. SK, VJ, AW and MHF developed the questionnaire. SK, VJ, AW and MHF were responsible for conducting the questionnaire. SK and VJ performed the descriptive analysis and the data management. SK, VJ and AW interpreted the data. SK, VJ and AW drafted the manuscript, while MHF revised it. All authors have read and approved the final manuscript and agreed to be accountable for all aspect of the work.

### **Disclosure**

MHF reports personal fees from DAK Gesundheit, outside the submitted work. The authors report no other conflicts of interest in this work.

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# Supplementary materials

 Table SI Questions on PSA testing V by general practitioners and urologists, n (%)

Question		GPs					Urologists	sts			
		Never	Rarely	Sometimes	Often	Always	Never	Rarely	Sometimes	Often	Always
On which occasions do you inform	Within an early cancer	(8) 8.7	7.3 (3)	9.8 (4)	19.5 (8)	56.1 (23)	0.0) 0	0.0) 0	0 (0:0)	1 (7.1)	13 (92.9)
your patient on PSA testing?	detection examination										
	In case of a positive	4 (9.8)	2 (4.9)	4 (9.8)	9 (22.0)	22 (53.7)	0.0) 0	0.0) 0	1 (7.1)	0.0) 0	13 (92.9)
	family anamnesis										
	In case of discomfort	7 (17.1)	9 (22.0)	8 (19.5)	8 (19.5)	9 (22.0)	0.0) 0	l (7.1)	2 (14.3)	8 (57.1)	3 (21.4)
	in the lower urinary										
	tract										
	In case of unclear	5 (12.2)	12 (29.3)	10 (24.4)	10 (24.4)	4 (9.8)	0.0) 0	I (7.I)	3 (21.4)	7 (50.0)	3 (21.4)
	discomfort										
How often do you discuss the fol-	Impact on overall	7 (17.1)	7 (17.1)	4 (9.8)	9 (22.0)	14 (34.1)	I (7.I)	I (7.I)	3 (21.4)	5 (35.6)	4 (28.6)
lowing aspects with your patients	mortality										
before performing a PSA test?	Impact on disease-	3 (7.3)	4 (9.8)	3 (7.3)	14 (34.1)	17 (41.5)	I (7.I)	I (7.I)	1 (7.1)	6 (42.9)	5 (35.7)
	specific mortality										
	Impact on risk of	5 (12.2)	4 (9.8)	8 (19.5)	9 (22.0)	15 (36.6)	0.0) 0	l (7.1)	1 (7.1)	9 (64.3)	3 (21.4)
	metastasis										
	Potential overdiagnosis	1 (2.4)	2 (4.9)	5 (12.2)	10 (24.4)	23 (56.1)	I (7.I)	I (7.I)	1 (7.1)	5 (35.7)	6 (42.9)
	Issue of false positives	0.0) 0	0.0)	6 (14.6)	12 (29.3)	23 (56.1)	0.0) 0	0.0) 0	0.0)	4 (28.6)	10 (71.4)
	Potential anxiety dur-	6 (14.6)	7 (17.1)	6 (14.6)	7 (17.1)	15 (36.6)	2 (14.3)	4 (28.6)	1 (7.1)	6 (42.9)	I (7.I)
	ing waiting on test										
	result										
	Potential follow-up	1 (2.4)	0.0)	8 (19.5)	11 (26.8)	21 (51.2)	0.0) 0	0.0) 0	1 (7.1)	3 (21.4)	10 (71.4)
	examinations if the										
	test result is										
	conspicuous										
	Adverse effects of the	1 (2.4)	4 (9.8)	12 (29.3)	11 (26.8)	13 (31.7)	0.0 (0)	0.0 (0)	6 (42.9)	5 (35.7)	3 (21.4)
	treatment										

Abbreviations: GPs, general practitioners; PSA, prostate-specific antigen.

Table S2 Questions on digital rectal examining by general practitioners and urologists, n (%)

Question		GPs					Urologists	sts			
		Never	Rarely	Sometimes	Often	Always	Never	Rarely	Always Never Rarely Sometimes	Often	Always
How often do you examine	During an early	0.0) 0	0.0) 0	4 (9.8)	4 (9.8)	33 (80.5)	0.0) 0	33 (80.5) 0 (0.0) 0 (0.0)	0.0)	2 (14.3)	12 (85.7)
digito rectally in the following											
situations?	examination										
	If there is blood in	0.0) 0	0.0) 0	3 (7.3)	13 (31.7)	25 (61.0) 0 (0.0)	0.0)	0.0) 0	0.0)	3 (21.4)	11 (78.6)
	the patient's stool										
	If the patient has	1 (2.4)	3 (7.3)	17 (41.5)	12 (29.3)	8 (19.5) 0 (0.0)	0.0)	0.0) 0	0.0)	4 (28.6)	10 (71.4)
	a voiding disorder										
	If the patient is	17 (41.5)	12 (29.3)	8 (19.5)	4 (9.8)	0.0) 0	0.0) 0 (0.0) 0	0.0) 0	4 (28.6)	6 (42.9)	4 (28.6)
	asymptomatic										

Abbreviation: GPs, general practitioners.

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