REVIEW





Is there any difference in urinary continence between bilateral and unilateral nerve sparing during radical prostatectomy? A systematic review and meta-analysis

Peng Xiang¹, Zhen Du¹, Di Guan¹, Wei Yan¹, Mingdong Wang¹, Danyang Guo¹, Dan Liu¹, Yuexin Liu¹ and Hao Ping^{1*}

Abstract

Context In men with prostate cancer, urinary incontinence is one of the most common long-term side effects of radical prostatectomy (RP). The recovery of urinary continence in patients is positively influenced by preserving the integrity of the neurovascular bundles (NVBs). However, it is still unclear if bilateral nerve sparing (BNS) is superior to unilateral nerve sparing (UNS) in terms of post-RP urinary continence. The aim of this study is to systematically compare the differences in post-RP urinary continence outcomes between BNS and UNS.

Methods The electronic databases of PubMed and Web of Science were comprehensively searched. The search period was up to May 31, 2023. English language articles comparing urinary continence outcomes of patients undergoing BNS and UNS radical prostatectomy were included. Meta-analyses were performed to calculate pooled relative risk (RR) estimates with 95% confidence intervals for urinary continence in BNS and UNS groups at selected follow-up intervals using a random-effects model. Sensitivity analyses were performed in prospective studies and robotic-assisted RP studies.

Results A meta-analysis was conducted using data from 26,961 participants in fifty-seven studies. A meta-analysis demonstrated that BNS improved the urinary continence rate compared to UNS at all selected follow-up points. RRs were 1.36 (1.14–1.63; p=0.0007) at ≤ 1.5 months (mo), 1.28 (1.08–1.51; p=0.005) at 3–4 mo, 1.12 (1.03–1.22; p=0.01) at 6 mo, 1.08 (1.05–1.12; p < 0.00001) at 12 mo, and 1.07 (1.00–1.13; p=0.03) at ≥ 24 mo, respectively. With the extension of the follow-up time, RRs decreased from 1.36 to 1.07, showing a gradual downward trend. Pooled estimates were largely heterogeneous. Similar findings were obtained through sensitivity analyses of prospective studies and robotic-assisted RP studies.

Conclusion The findings of this meta-analysis demonstrate that BNS yields superior outcomes in terms of urinary continence compared to UNS, with these advantages being sustained for a minimum duration of 24 months. It may be due to the real effect of saving the nerves involved. Future high-quality studies are needed to confirm these findings.

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Keywords Radical prostatectomy, Urinary continence, Bilateral nerve sparing, Unilateral nerve sparing, Systematic review

Introduction

Globally, prostate cancer (PCa) is the second most common cancer and the fifth leading cause of cancer death among men in 2020, with approximately 1,414,259 new cases and 375,000 deaths [1, 2]. Radical prostatectomy (RP) is one of the gold standard treatments for patients diagnosed with localized or locally advanced prostate cancer. It is widely acknowledged that urinary incontinence and erectile dysfunction are significant causes of low quality of life in many surgical men [2, 3]. At 12 months (mo), 16% of post-RP men are incontinent (using a no-pad definition) [4]. The reported potency rates following robot-assisted RP are highly variable, ranging from 54-90% [5]. Long-term urinary incontinence has been linked to a number of complex issues in individuals with prostate cancer, such as obsessivecompulsive about restroom locations and preventing leaks, as well as feelings of shame, helplessness, and uncleanliness when control is compromised [6].

In RP, different surgical approaches (open, laparoscopic, or robotic-assisted) are used, but postoperative continence is mainly determined by surgical technique, including preservation techniques and reconstructing techniques [2, 7]. In general, the retention of various structures like the neurovascular bundles (NVBs) and bladder neck aids in the control of urination [8]. The preservation of NVBs during RP has been shown to lead to an earlier return of continence [6, 9]. Recent meta-analyses reported that patients who had any nerve sparing (NS) surgery (i.e., bilateral nerve sparing, unilateral nerve sparing, or unspecified) had significantly better continence outcomes compared to those who had non-nerve sparing surgery [3, 6, 9]. Furthermore, Reeves et al. [6] demonstrated that there was only a statistically significant difference in continence outcomes between bilateral nerve sparing (BNS) and unilateral nerve sparing (UNS) at short-term follow-up (≤ 1.5 mo). However, Nguyen et al. [3] indicated that lower rates of urinary incontinence were significantly observed with BNS compared to UNS at 1 year. If sparing two-sided NVBs has a real advantage in postoperative urinary continence, then preservation of continence should be an independent indication for BNS, which will be crucial for clinical practice. Until now, data concerning bilateral vs. unilateral nerve-sparing radical prostatectomy in urinary continence has been widely reported and the results are controversial.

In this research, we aimed to conduct a systematic review and meta-analysis to assess whether there were differences in urinary continence outcomes after RP surgery between BNS and UNS in both short-term and long-term follow-up. The hypothesis posits that adopting a BNS approach may mitigate the incidence of urinary incontinence.

Methods

A systematic review and meta-analysis was conducted in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-analysis (PRISMA) statement. The protocol for this study was registered with PROS-PERO (CRD42022378340, https://www.crd.york.ac.uk/ PROSPERO/#recordDetails).

Search strategy and eligibility criteria

To identify potentially relevant studies, searches were conducted in the PubMed and Web of Science electronic databases on May 31, 2023. Based on the absence of a formal description of NS surgery before 1982, the search was limited to studies published after that time.

We included studies reporting the outcomes of urinary continence in men treated for PCa with a BNS RP (intervention) and a UNS RP (control). Specifically, this review included retrospective and prospective studies that evaluated the comparison of urinary incontinence outcomes between UNS and BNS. Cross-sectional studies and observational studies without a control group (i.e., singlecohort studies) were not included. The present study did not attempt to analyze more specific or alternative types of nerve sparing, such as intra/interfascial versus standard, risk-stratified NS, or sural nerve grafting. We did not exclude studies based on surgical approaches such as open RP (ORP), laparoscopic RP (LRP), or roboticassisted RP (RARP). In the search, the following terms were used: ["nerve sparing"] AND ["prostatectomy"] AND ["unilateral "OR "bilateral"]. The above keywords are searched using "all fields". See Additional files 1 and 2 for detailed search strategies.

A meta-analysis of relevant prospective and retrospective studies with sufficient data was conducted. The scope of the review according to the PICO process (Patient, Intervention, Comparison, Outcomes) is as follows: P-patients with prostate cancer; I- radical prostatectomy with BNS; C-radical prostatectomy with UNS; O-urinary continence outcomes. The search was limited to Englishlanguage publications. To identify additional potentially relevant studies, we manually searched the reference lists of relevant publications and reviews. When data is duplicated, more recent or comprehensive studies are preferred.

Outcome

Postoperative urinary continence was the primary outcome of this systematic review and meta-analysis. Continence rates from included studies were pooled in this meta-analysis. We investigate the effect of BNS versus UNS on continence rates at selected follow-up intervals including \leq 1.5 mo, 3–4 mo, 6 mo, 12 mo, and \geq 24 mo.

Study selection and data extraction

Two authors screened the search results (titles and abstracts), and any disagreements were resolved by consensus. In line with the previously outlined selection criteria, the full texts of all potentially relevant publications were retrieved for review.

Data extracted involved patient age, sample size, continence definition, and surgical approach. To conduct the meta-analysis, the total number of participants and events were extracted (defined as a number of continent men). Raw numbers can also be calculated based on hazard ratios, relative risks, or odds ratios results. Any differences of opinion regarding data extraction were resolved through discussion and, if necessary, consultation with a third author to reach an agreement.

Synthesis of results and statistical analysis

The risk of bias in each study was independently evaluated by two authors according to the method published in the study of Reeves et al. [6]. This evaluation covered baseline continence, outcome assessment (the definition of urinary continence in each paper), comparability of groups (comparison of clinical information including age, tumor stage, and Gleason classification), NS assessment (details of NS surgical procedures), surgical technique variations (additional surgical procedures, such as puboprostatic ligaments preservation, bladder neck reconstruction and posterior rhabdosphincter reconstruction) and other issues like unexplained loss to follow-up and selective outcome reporting. Each comparison was measured as a risk ratio (RR) with 95% confidence interval. Recovery of continence is more likely in the intervention group if the RR > 1.0.

Results of each study were grouped by type of NS (BNS, or UNS), as well as stratified by the timing of outcome reporting. Based on all available results in included studies, the outcome timing categories are $\leq 1.5 \text{ mo}$, 3-4 mo, 6 mo, 12 mo, and $\geq 24 \text{ mo}$. A Mantel-Haenszel random-effects model was used to calculate pooled.

RRs for each time category. The I² statistic (I² \geq 50%) and chi-square test ($p \leq 0.10$) were used to assess heterogeneity.

Sensitivity analysis was conducted for studies that used a prospective design or prospectively collected data. At the same time, a sensitivity analysis of RARP studies was also conducted. Moreover, due to few studies, it is unlikely that separate sub-analyses could be performed based on the type of procedure (e.g., robotic-assisted RP versus other RP) and urinary continence definition (e.g., no pad vs. other).

To assess the risk of publication bias, funnel plots and the Egger test of funnel plot symmetry were generated for all primary outcomes. The Egger test is based on linear regression of the intervention effect estimate against its standard error weighted by the inverse of the intervention effect estimate's variance. Examine potential publication bias through visual examination of funnel plots. Importantly, the significant publication bias is indicated by a p-value < 0.05. RevMan5.3 and STATA 12 software were used to perform all statistical analyses and generate forest plots.

Results

Study selection and characteristics

Our search yielded 1249 unique records. Ultimately, fiftyseven studies were included for quantitative synthesis. A total of forty-six prospective longitudinal cohort studies [10–55] and eleven retrospective cohort studies [56–66] were included in this review (Table 1). The study sample size ranged from 15 to 2019 participants, with a total of 26,961 patients included in the final analysis. A series of studies were conducted between 1997 and 2023. Most patients included in the studies were undergoing RARP. Some studies used open surgery (retropubic or perineal) or extraperitoneal laparoscopic surgery. About 90% of the research originated from Europe and the United States. The demographic data of fifty-seven patient-series were comparable, with an average age range of 51-69 years and an average follow-up range of immediately after catheter removal to >60 mo. The most common definition of urinary continence in included studies was no pad use.

Assessment of methodology of included studies

Because the BNS/UNS group stratification characteristics were not fully described in most of the included studies, the baseline imbalance cannot be determined. In the case of the data provided, selection bias was evident. Patients who underwent the BNS procedure were more likely to have younger ages, lower Gleason scores, better baseline sexual function, and a favorable clinical stage. A few studies also adjusted for age, comorbidities,

Table 1 Characteristics of the included studies

itudy, year	Sample size	Surgical approach Age, yr, mea median		Timing of outcome, mo	Continence definition	
Prospective						
Albayrak 2010 [10]	BNS 73, UNS 12	Perineal	62	3	No pad	
Asimakopoulos 2019 [11]	BNS 69, UNS 10	RARP	65	Immediate	No pad	
Avulova 2018 [12]	BNS 805, UNS 186	80% RARP, 19% ORP, 1% other	19% ORP, BNS 61, UNS 63 36		No pad	
Berg 2014 [13]	BNS 85, UNS 72	RARP	BNS 60, UNS 63	3, 6, 12, 24	No pad	
Berry 2009 [14]	BNS 341, UNS 89 65%RARP, 12% LRP, BNS 59, UNS 59 36 23% ORP		36	Return to 75% of base line continence score (UCLA-PCI)		
Bhat 2022 [15]	BNS 1308, UNS 532	RARP	51	12	No pad	
Budaus 2009 [16]	BNS 464, UNS 173	Retropubic	63	12	0–1 pad per day	
Burkhard 2006 [17]	BNS 75, UNS 322	Retropubic	64	12	No pad	
Choi 2011 [18]	BNS 469, UNS 89	RARP	BNS 58, UNS 59 4, 12, 24		No pad	
Collette 2021 [19]	BNS 990, UNS 466	RARP	66	12	0–1 pad per day	
d'Altilia 2022 [<mark>20</mark>]	BNS 120, UNS 49	59%RARP; 41% ORP	66	3, 6, 12	24-h pad test ≤ 20g/d	
Dalkin 2006 [21]	BNS 53, UNS 68	Retropubic	63	12, 24	No pad	
El-Hakim 2015 [22]	BNS 167, UNS 28	RARP	60	3	No pad	
Feng 2020 [23]	BNS 187, UNS 84	RARP	62	15	No pad	
Fossati 2017 [24]	BNS 1351, UNS 144	RARP	63	12	No pad	
Geraerts 2013 [25]	BNS 112, UNS 44	36%RARP, 64%ORP	62	12	0 g urine leakage	
Hatiboglu 2015 [26]	BNS 697, UNS 202	57%RARP, 43%Retro- pubic			No pad	
Hinata 2014 [27]	BNS 35, UNS 92	RARP	BNS 62, UNS 63	1, 3, 6	No pad	
Holze 2019 [28]	BNS 153, UNS 57	46.2%RARP, 53.8%ORP	65	3	No pad	
Kim 2019 [29]	BNS 285, UNS 214	RARP	65	1, 3, 6, 12	No pad and no leakag	
Ko 2012 [<mark>30</mark>]	BNS 779, UNS 394	RARP	60	1.5, 3, 6	No pad and no leakag	
Kohjimoto 2022 [31]	BNS 92, UNS 199	RARP	69	1, 3, 6, 12, 24	No pad	
Kováčik 2019 [<mark>32</mark>]	BNS 60, UNS 28	RARP	64	0.5, 3, 6, 12	0–1 pad per day	
Kowalczyk 2013 [33]	BNS 490, UNS 120	RARP	60	5, 12	No pad	
Kung 2015 [34]	BNS 33, UNS 10	Retropubic	60	57	No pad	
Lavigueur-Blouin 2015 [35]	BNS 201, UNS 52	RARP	60	1	No pad	
Lee 2010 [36]	BNS 58, UNS 15	RARP	59	1.5	No pad	
Marien 2008 [37]	BNS 538, UNS 72	Retropubic	BNS 57, UNS 59	24	Total control or occa- sional leakage	
Nandipati 2007 [<mark>38</mark>]	BNS 66, UNS 25	ORP	BNS 62, UNS 64	3, 6, 12, 24, >60	No pad	
Novara 2010 [39]	BNS 201, UNS 22	RARP	62	12	No leakage	
Nyarangi-Dix 2020 [40]	BNS 156, UNS 86	RARP	65	12, 24	No pad	
Pagliarulo 2020 [41]	BNS 163, UNS 68	LRP	65	12	No pad	
Pick 2011 [42]	BNS 357, UNS 143	RARP	BNS 60, UNS 63	1, 3, 12	No pad	
Reichert 2022 [43]	BNS 25, UNS 27	RARP	64	12	No pad	
Rigatti 2012 [44]	BNS 24, UNS 9	RARP	66	1, 3	No leakage	
Sammon 2013 [45]	BNS 1015, UNS 125	RARP	60	Immediate	No pad	
Scarcia 2018 [46]	BNS 208, UNS 201	RARP	65	1, 3, 12	0–1 pad per day	
Steineck 2015 [47]	BNS 970, UNS 959	78%RARP, 22%ORP	63	12	No pad	
Suardi 2012 [48]	BNS 900, UNS 49	Retropubic	64	12, 24	No pad	
Talcott 1997 [49]	BNS 28, UNS 38	ORP	BNS 61, UNS 62	3, 12	No pad	
Theissen 2019 [50]	BNS 76, UNS 23	50%RARP, 50%ORP	66	Immediate	Urine loss (≤ 10g) within 1h	
Toren 2009 [51]	BNS 159, UNS 32	ORP	BNS 59, UNS 60	12	No or rare urine leakag	

Study, year	dy, year Sample size		Age, yr, mean or median	Timing of outcome, mo	Continence definition	
Tsikis 2017 [52]	BNS 396, UNS 51	RARP	58	12	No pad	
Tzou 2009 [53]	BNS 73, UNS 112	Retropubic	63	12, 24	No pad	
Van der Poel 2009 [54]	BNS 61, UNS 72	RARP	60	6	No involuntary urine loss	
Van der Slot 2023 [55]	BNS 340, UNS 202	RARP	68	6, 12, 24	0–1 pad per day	
Retrospective						
Chung 2020 [<mark>56</mark>]	BNS 125, UNS 72	70%RARP, 30%ORP	68	12	No pad	
Fosså 2019 [57]	BNS 165, UNS 242	RARP	62	24	No pad	
Greco 2011 [58]	BNS 250, UNS 207	Extraperitoneal LRP	BNS 59, UNS 60	1, 3, 12	No pad	
Hinata 2019 [59]	BNS 46, UNS 137	RARP	BNS 65, UNS 66	24	No pad	
Kadono 2015 [<mark>60</mark>]	BNS 15, UNS 36	RARP	BNS 64, UNS 65	12	24-h pad test≤2g/day	
Lee 2013 [61]	BNS 100, UNS 54	Extraperitoneal LRP	66	3	No pad	
Noël 2022 [<mark>62</mark>]	BNS 391, UNS 138	RARP	57	1.5	No leakage	
Palisaar 2015 [63]	BNS 1332, UNS 687	RARP or ORP	64	1.5	0–1 pad per day	
Punnen 2014 [64]	BNS 157, UNS 74	RARP	NA	6	No pad	
Shikanov 2011 [65]	BNS 1021, UNS 322	RARP	60	12	No pad	
Wang 2014 [66]	BNS 2, UNS 13	RARP	64	12	No pad	

Table 1 (continued)

BNS bilateral nerve sparing, UNS unilateral nerve sparing, RARP robot-assisted radical prostatectomy, LRP laparoscopic radical prostatectomy, NA not available, ORP open radical prostatectomy, UCLA-PC/University of California, Los Angeles, Prostate Cancer Index

and history of lower abdominal surgery when calculating RRs. Indeed, urinary continence outcomes may be influenced by various confounding factors, including age, body mass index, prior surgical history, prostate volume, membranous urethral length, tumor staging and grading, and surgeon experience. However, despite the lack of thorough characterization of NVB preservation, even in patients with low-risk prostate cancer, its implementation was associated with a higher risk of cancer at the margins of the excised tissue [67].

Baseline continence was seldom reported. In several studies, there was no urinary incontinence reported before surgery. Several other studies have reported similar baseline continence scores in the comparison groups.

Few studies have described how to define or record the NS status. In one study, it is stated that >70% of the bundles preserved in situ can be regarded as NS [18]. Several studies describe retrospective reviews of surgical reports to determine NS status. Some studies directly describe NS using interfascial, extrafascial, or intrafascial techniques. At the same time, some of the studies were conducted concurrently with the preservation of the puboprostatic ligament or bladder neck reconstruction.

The assessment of results was variable. Most studies used the definition of no-pad for continence. Few studies used validated instruments to determine incontinence status. Even when a validated tool was used, it was often unclear who submitted the questionnaire and whether the assessor was blinded. Potential selective outcome reporting or unexplained loss to follow-up was apparent in many studies (Additional file 3).

Urinary continence outcomes

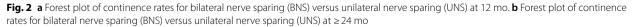
Patients who underwent the BNS procedure had significantly better continence outcomes compared to those who underwent the UNS procedure. RRs were 1.36 (1.14–1.63; p=0.0007) at ≤ 1.5 mo, 1.28 (1.08–1.51; p=0.005) at 3–4 mo, 1.12 (1.03–1.22; p=0.01) at 6 mo, 1.08 (1.05–1.12; p<0.00001) at 12 mo and 1.07 (1.00–1.13; p=0.03) at ≥ 24 mo. Notably, the RRs decreased gradually as the follow-up time extended. Figures 1 and 2 showed the meta-analysis of urinary continence outcomes for BUS compared to UNS at selected follow-up points.

Sensitivity analysis revealed consistent overall results when prospective studies or RARP studies were considered alone. The RRs of BNS compared to UNS in prospective studies at $\leq 1.5 \text{ mo}$, 3-4 mo, 6 mo, 12 mo and $\geq 24 \text{ mo}$ were 1.47 (1.05-2.04; p=0.02), 1.33 (1.07-1.66; p=0.01), 1.12 (1.02-1.23; p=0.01), 1.09 (1.04-1.13; p<0.0001) and 1.07 (1.00-1.14; p=0.06), respectively (Additional files 4 and 5). The RRs of BNS compared to UNS in RARP patients at $\leq 1.5 \text{ mo}$, 3-4 mo, 6 mo, 12 mo and $\geq 24 \text{ mo}$ were 1.47 (1.06-2.03; p=0.02), 1.35 (1.00-1.81; p=0.05), 1.14 (1.03-1.26; p=0.01), 1.11 (1.05-1.16; p<0.0001) and 1.09 (1.00-1.18; p=0.04), respectively (Additional files 6 and 7).

Study or Subgroup	BNS		UN		Woight	Risk Ratio M-H, Random, 95% Cl	Risk Ratio M-H. Random, 95% Cl
					-		
Asimakopoulos 2019	30	69	2	10	1.6%	2.17 [0.61, 7.73]	-
Greco 2011	107	250	66	207	7.7%	1.34 [1.05, 1.72]	
latiboglu 2015	125	697	33	202	6.7%	1.10 [0.77, 1.56]	
linata 2014	19	35	46	92	6.5%	1.09 [0.75, 1.57]	
(im 2019	78	285	33	214	6.5%	1.77 [1.23, 2.56]	
Ko 2012	552	779	292	394	8.9%	0.96 [0.89, 1.03]	1
Kohjimoto 2022	12	92	7	199	2.7%	3.71 [1.51, 9.11]	
kováčik 2019	15	60	12	28	4.4%	0.58 [0.32, 1.08]	
avigueur-Blouin 2015.	98	201	18	52	6.2%	1.41 [0.94, 2.10]	-
.ee 2010	33	58	6	15	4.0%	1.42 [0.74, 2.75]	
loël 2022	210	391	64	138	8.1%	1.16 [0.95, 1.42]	-
Palisaar 2015	973	1332	456	687	8.9%	1.10 [1.03, 1.17]	
Pick 2011	117	357	43	143	7.3%	1.09 [0.81, 1.46]	+
Rigatti 2012	14	24	3		2.4%	1.75 [0.65, 4.68]	
Sammon 2013	184	1015	22	125	6.2%	1.03 [0.69, 1.54]	<u> </u>
Scarcia 2018	169	208	26	201	6.5%	and the second formation of the second form	-
Theissen 2019	48	76	10	201		6.28 [4.36, 9.04]	
neissen 2019	40	70	10	23	5.3%	1.45 [0.88, 2.39]	
Total (95% CI)		5929		2739	100.0%	1.36 [1.14, 1.63]	◆
Total events	2784		1139				
Heterogeneity: Tau ² = 0.	.09; Chi ² =	145.22	2, df = 16	(P < 0.	00001); l²	= 89%	0.01 0.1 1 10 10
Test for overall effect: Z	= 3.41 (P	= 0.000	07)				Favours [UNS] Favours [BNS]
)	BNS		UNS			Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% C	M-H, Random, 95% Cl
Albayrak 2010	58	73	7	12	4.6%	1.36 [0.83, 2.23]	+
Berg 2014	34	85	17	72	4.6%	1.69 [1.04, 2.77]	— —
Choi 2011	178	377	35	72	6.3%	0.97 [0.75, 1.26]	+
			37		6.9%	1.11 [0.93, 1.33]	-
d'Altilia N 2022	101	120		49			
El-Hakim 2015	83	167	13	28	5.1%	1.07 [0.70, 1.64]	
Greco 2011	220	250	153	207	7.3%	1.19 [1.08, 1.31]	
Hinata 2014	24	35	59	92	6.2%	1.07 [0.82, 1.40]	Ť
Holze 2019	87	153	24	57	5.8%	1.35 [0.97, 1.89]	
Kim 2019	142	285	75	214	6.6%	1.42 [1.14, 1.77]	-
Ko 2012	684	784	350	395	7.4%	0.98 [0.94, 1.03]	•
Kohjimoto 2022	25	92	27	199	4.6%	2.00 [1.23, 3.25]	
							-
Kováčik 2019	36	60	18	28	5.7%	0.93 [0.66, 1.32]	_
_ee 2013	40	100	27	54	5.6%	0.80 [0.56, 1.15]	
Vandipati 2007	39	66	14	25	5.2%	1.06 [0.71, 1.58]	T
Pick 2011	233	355	91	141	7.1%	1.02 [0.88, 1.17]	Ť
Rigatti 2012	5	10	2	6	1.4%	1.50 [0.41, 5.45]	
Scarcia 2018	194	208	45	201	6.3%	4.17 [3.21, 5.40]	-
Talcott 1997	14	28	9	38	3.4%	2.11 [1.07, 4.17]	
Fotal (95% CI)		3248		1890	100.0%	1.28 [1.08, 1.51]	◆
Fotal events	2197		1003				
Heterogeneity: Tau ² = 0				7 (P < 0	0.00001);	l ² = 92%	0.01 0.1 1 10 10
Test for overall effect: Z		r = 0.00					Favours [UNS] Favours [BNS]
	BNS	-	UNS			Risk Ratio	Risk Ratio
Study or Subgroup					vord Terrarent	M-H, Random, 95% C	M-H, Random, 95% Cl
Berg 2014	63	85	42	72	7.3%	1.27 [1.01, 1.60]	
d'Altilia N 2022	101	120	37	49	9.3%	1.11 [0.93, 1.33]	T.
Hinata 2014	28	35	72	92	8.5%	1.02 [0.84, 1.25]	+
Kim 2019	210	280	139	214	11.7%	1.15 [1.02, 1.30]	-
(0.2012	552	779	292	394	13.5%	0.96 [0.89, 1.03]	•
Ko 2012	57	92	75	199	7.0%	1.64 [1.29, 2.09]	-
		60	23	28	8.7%	1.10 [0.90, 1.33]	+
Kohjimoto 2022		490					_ _
Kohjimoto 2022 Kováčik 2019	54 77		16	120	2.5% 6.5%	1.18 [0.71, 1.94]	_
Kohjimoto 2022 Kováčik 2019 Kowalczyk 2013	77		40		h 5%	1.02 [0.79, 1.31]	
Kohjimoto 2022 Kováčik 2019 Kowalczyk 2013 Nandipati 2007	77 51	66	19	25			
Kohjimoto 2022 Kováčik 2019 Kowalczyk 2013 Nandipati 2007 Punnen 2014	77 51 89	66 157	36	74	6.1%	1.17 [0.89, 1.53]	T
Kohjimoto 2022 Kováčik 2019 Kowalczyk 2013 Nandipati 2007 Punnen 2014 /an der Poel 2009	77 51 89 41	66 157 61	36 34	74 72	6.1% 5.4%	1.17 [0.89, 1.53] 1.42 [1.05, 1.92]	
Kohjimoto 2022 Kováčik 2019 Kowalczyk 2013 Nandipati 2007 Punnen 2014	77 51 89	66 157	36	74	6.1%	1.17 [0.89, 1.53]	
Kohjimoto 2022 Kováčik 2019 Kowalczyk 2013 Vandipati 2007 Punnen 2014 /an der Poel 2009 /an der Slot 2023	77 51 89 41 198	66 157 61 225	36 34	74 72 144	6.1% 5.4% 13.5%	1.17 [0.89, 1.53] 1.42 [1.05, 1.92] 0.99 [0.92, 1.07]	
Kohjimoto 2022 Kováčik 2019 Kowalczyk 2013 Vandipati 2007 Punnen 2014 /an der Poel 2009 /an der Slot 2023 Fotal (95% CI)	77 51 89 41 198	66 157 61	36 34 128	74 72 144	6.1% 5.4%	1.17 [0.89, 1.53] 1.42 [1.05, 1.92]	
Kohjimoto 2022 Kováčik 2019 Kowalczyk 2013 Vandipati 2007 Punnen 2014 /an der Poel 2009 /an der Slot 2023	77 51 89 41 198	66 157 61 225 2450	36 34 128 913	74 72 144 1483	6.1% 5.4% 13.5% 100.0%	1.17 [0.89, 1.53] 1.42 [1.05, 1.92] 0.99 [0.92, 1.07] 1.12 [1.03, 1.22]	

Fig. 1 a Forest plot of continence rates for bilateral nerve sparing (BNS) versus unilateral nerve sparing (UNS) at \leq 1.5 mo. b Forest plot of continence rates for bilateral nerve sparing (BNS) versus unilateral nerve sparing (UNS) at 3–4 mo. c Forest plot of continence rates for bilateral nerve sparing (BNS) versus unilateral nerve sparing (UNS) at 3–4 mo. c Forest plot of continence rates for bilateral nerve sparing (UNS) at 6 mo

а	BNS	5	UNS	5		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total			Weight	M-H. Random, 95% C	
Berg 2014	73	85	55	72	2.6%	1.12 [0.96, 1.31]	-
Bhat 2022	453	1308	160	532	2.7%	1.15 [0.99, 1.34]	+
Budäus 2009	439	464	160	173	5.1%	1.02 [0.98, 1.07]	•
Burkhard 2006	74	75	311	322	5.4%	1.02 [0.99, 1.06]	•
Choi 2011	256	303	38	52	2.3%	1.16 [0.97, 1.37]	-
Chung 2020	109	125	59	72	3.2%	1.06 [0.94, 1.21]	+
Collette 2021	841	990	332	466	4.7%	1.19 [1.12, 1.27]	-
d'Altilia N 2022	110	120	40	49	2.8%	1.12 [0.97, 1.30]	-
Dalkin 2006	44	53	54	68	2.3%	1.05 [0.88, 1.24]	+
Feng 2020	136	187	58	84	2.4%	1.05 [0.89, 1.25]	+
Fossati 2017	984	1351	107	144	3.8%	0.98 [0.89, 1.08]	+
Geraerts 2013	104	112	39	44	3.4%	1.05 [0.93, 1.18]	+
Greco 2011	243	250	182	207	4.9%	1.11 [1.05, 1.17]	-
Kadono 2015	12	15	26	36	0.9%	1.11 [0.80, 1.53]	
Kim 2019	269	285	167	214	4.4%	1.21 [1.12, 1.31]	-
Kohjimoto 2022	63	92	107	199	2.1%	1.27 [1.05, 1.54]	-
Kováčik 2019	53	60	24	28	2.2%	1.03 [0.86, 1.23]	+
Kowalczyk 2013	98	490	10	120	0.3%	2.40 [1.29, 4.46]	
Nandipati 2007	57	490	23	25	2.7%	to to a firm of the coal	4
and the second s						0.94 [0.81, 1.09]	-
Novara 2010	180	201	19	22	2.3%	1.04 [0.87, 1.23]	-
Nyarangi-Dix 2020	116	156	57	86	2.2%	1.12 [0.94, 1.34]	
Pagliarulo 2020	152	163	62	68	4.2%	1.02 [0.94, 1.11]	I I I I I I I I I I I I I I I I I I I
Pick 2011	296	332	120	135	4.6%	1.00 [0.93, 1.08]	
Reichert 2022	21	25	11	27	0.5%	2.06 [1.27, 3.35]	
Scarcia 2018	204	208	174	201	4.9%	1.13 [1.07, 1.20]	
Shikanov 2011	726	1021	215	322	4.1%	1.06 [0.98, 1.16]	[
Steineck 2015	837	970	716	959	5.2%	1.16 [1.11, 1.21]	•
Suardi 2012	716	900	31	49	1.7%	1.26 [1.01, 1.56]	
Talcott 1997	16	28	17	38	0.5%	1.28 [0.79, 2.06]	
Toren 2009	116	159	27	32	2.2%	0.86 [0.72, 1.03]	
Tsikis 2017	261	396	29	51	1.4%	1.16 [0.90, 1.49]	-
Tzou 2009	62	73	86	112	2.9%	1.11 [0.96, 1.27]	T
Van der Slot 2023	206	223	121	130	4.8%	0.99 [0.93, 1.05]	t and the second s
Wang 2014	2	2	12	13	0.4%	0.93 [0.55, 1.60]	
Total (05% CI)		11288		5152	100.0%	1.08 [1.05, 1.12]	
Total (95% CI)	0000	11200	2640	5152	100.076	1.00 [1.05, 1.12]	
Total events	8329	- 445 4	3649		00004)	2 - 740/	
Heterogeneity: Tau ² = 0				3 (P < 0	0.00001); 1	r = 71%	0.01 0.1 1 10 100
Test for overall effect: 2	2 = 4.01 (i	2 < 0.00	001)				Favours [UNS] Favours [BNS]
b	BNS	6	UNS	5		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% C	I M-H. Random, 95% CI
Avulova 2018	676	805	146	186	8.4%	1.07 [0.99, 1.16]	•
Berg 2014	73	75	58	72	7.1%	1.21 [1.07, 1.36]	-
Berry 2009	187	341	40	89	3.6%	1.22 [0.95, 1.57]	
Choi 2011	138	146	25	27	7.3%	1.02 [0.91, 1.14]	+
Dalkin 2006	22	23	45	53	6.3%	1.13 [0.98, 1.30]	+
Fosså 2019	135	165	179	242	7.6%	1.11 [1.00, 1.23]	-
Hinata 2019	35	46	104	137	5.0%	1.00 [0.83, 1.21]	+
Kohjimoto 2022	70	92	116	199	5.6%	1.31 [1.11, 1.54]	-
Kung 2015	24	33	9	10	2.9%	0.81 [0.60, 1.08]	
Marien 2008	516	538	71	72	9.7%	0.97 [0.94, 1.00]	-
Nandipati 2007	62	66	24	25	7.7%	0.98 [0.88, 1.08]	+
Nyarangi-Dix 2020	131	156	68	86	6.8%	1.06 [0.93, 1.21]	÷
Suardi 2012		900	37		5.7%		
Tzou 2009	756			49		1.11 [0.95, 1.31]	Ļ
	55	59	70	82	7.3%	1.09 [0.98, 1.22]	ſ
Van der Slot 2023	118	123	79	82	9.2%	1.00 [0.94, 1.05]	
Total (95% CI)		3568		1411	100.0%	1.07 [1.00, 1.13]	
Total events	2998		1071				
Heterogeneity: Tau ² =		= 68.63		(P < 0.	.00001): l ²	= 80%	
Test for overall effect:							0.01 0.1 1 10 100 Favours [UNS] Favours [BNS]
		<i>.</i> .					



Heterogeneity and publication bias

Clinical and methodological inconsistencies were apparent in all included studies. There were significant differences in participant characteristics, including baseline urinary continence and age. Variations in surgical technique, such as preserving of the bladder neck, sparing of puboprostatic ligament, and posterior reconstruction, also varied from study to study or were not described in detail. Furthermore, some studies restricted surgery to high-volume surgeons or surgeons with a minimum level of expertise. In addition, there were also differences in the methods used to assess postoperative incontinence status. In general, high inter-study heterogeneity was observed at each time point of urinary continence. Consequently, a random-effects model was utilized in the meta-analyses. Statistical analysis of funnel plot asymmetry using Egger linear regression revealed no convincing evidence of publication bias, except for the outcome at 6 mo and ≥ 24 mo. Outcomes at 6 mo and ≥ 24 mo showed publication bias, possibly due to a limited number of included studies. P values for publication bias were 0.06, 0.06, 0.01, 0.21, and 0.01 at \leq 1.5mo, 3–4 mo, 6mo, 12mo, and ≥ 24 mo, respectively (Additional file 8).

Discussion

In this systematic review, we analyzed the differences in urinary continence between preserving bilateral NVBs and unilateral NVBs. The meta-analysis showed a correlation between BNS and improved urinary continence outcomes at all follow-up intervals, although this improvement gradually diminished with longer follow-up periods. Unlike the findings reported by Reeves et al. [6] and Nguyen et al. [3], which stated that BNS only exhibited good urinary continence at 1.5 months and 1 year after surgery, respectively.

The pathophysiology of post-RP urinary incontinence is not fully understood. Several factors are associated with the risk of postoperative urinary incontinence, including patient characteristics (e.g., body mass index, age, prostate volume, comorbidities) and provider-related factors (surgeon experience, skill, central volume, etc.) [68–70]. As far as the surgical methods are concerned, urethral sphincter preservation and nerve-sparing as well as the newly invented hood technique, can be effective in preventing post-RP urinary incontinence [70, 71]. Although most authors agree that the pudendal nerve innervates the rhabdosphincter, several anatomical studies have indicated abnormal intrapelvic somatic nerves to the sphincter [8]. Anatomic studies have also shown a partially intrapelvic route for the pudendal nerve branches that go on to innervate the urethral sphincter [7]. The impact of BNS on urinary continence outcomes may be explained by the preservation of these intrapelvic nerves to the rhabdosphincter. As the follow-up time prolongs, the decrease in the difference between BNS and UNS may be attributed to the compensation of other continence mechanisms, such as the pelvic floor musculature, bladder neck sparing, Retzius-sparing RARP and preservation of tissue around the urethra. The most common standard for BNS RP was the presence of low-risk disease [67]. The same but less stringent criteria were used to select patients for UNS: PSA < 10 ng/ml or GS ≤ 6 or normal DRE on the NS side, with or without biopsy core positive information [67]. The decision to perform BNS surgery in these individuals should be made on a case-by-case basis, considering risk stratification based on comprehensive clinical examination, biopsy findings, and imaging results. If BNS is appropriate from an oncologic standpoint, it should be taken into consideration as it might result in better potential for urinary continence following surgery. In addition, UNS can be selected for patients with intermediate- and high-risk diseases who need nerve preservation because its long-term urinary continence outcomes are comparable to those of BNS. Randomized controlled trials comparing BNS with UNS RP are unlikely to be designed for ethical reasons. In the future, to determine the best candidates for BNS RP and UNS RP, prospective multicenter trials with high methodological quality and long-term follow-up for patients with intermediate- and high-risk PCa are required. A deeper understanding of the risk factors for urinary continence may be achievable with sufficiently large sample sizes and multivariate analysis adjusted for specific confounders.

Even though this is the most thorough analysis comparing urinary continence outcomes of BNS to UNS during RP, various limitations must be taken into account when interpreting these data. This meta-analysis is compromised by the absence of randomized controlled trials. Prospective randomized controlled trials, however, are challenging to undertake because the nerve sparing technique may be modified during surgery based on the level of tumor involvement, intraoperative pathology, and other ethical considerations. If the intraoperative frozen pathological results indicate positive surgical margins when the patient undergoes planned BNS, the surgical method may be changed to UNS or NNS for the patient's ethical consideration. Moreover, our study did not include studies that were not published or in English. Although Egger's linear regression did not disclose any conclusive evidence of publication bias, assessing publication bias is inherently challenging when there are few studies included. There was significant heterogeneity among studies in terms of urinary continence, which we were unable to fully explain. Heterogeneity can be caused by a variety of factors, including age, prostate volume,

membranous urethral length, tumor staging and grading, surgeon experience, and variations in surgical technique. Many studies did not provide enough information to allow for adjustments, and stratification of research outcomes based on surgeon experience, patient age, or other factors was not done. Additionally, variations in the definition of NS status could affect the outcomes. Intrafascial, interfascial, or extrafascial nerve preservation surgery can result in inconsistencies in urinary continence outcomes.

Conclusion

This meta-analysis demonstrates that BNS results in superior urinary continence outcomes compared to UNS in all selected postoperative follow-up periods. This superiority persists for ≥ 24 mo. We speculate that the preservation of the intrapelvic nerves supplying the rhab-dosphincter may be the cause of this relationship. If BNS is deemed appropriate from an oncological perspective, it should be duly considered. High-quality cohort studies are recommended to corroborate the foregoing findings and additional research into the mechanisms of post-RP incontinence.

Supplementary Information

The online version contains supplementary material available at https://doi. org/10.1186/s12957-024-03340-6.

Supplementary Material 1.	
Supplementary Material 2.	
Supplementary Material 3.	
Supplementary Material 4.	
Supplementary Material 5.	
Supplementary Material 6.	
Supplementary Material 7.	
Supplementary Material 8.	

Authors' contributions

All authors contributed to the study conception and design. Material preparation was performed by P Xiang, Z Du, D Guan, M Wang, D Guo and W Yan. Data collection and analysis were performed by P Xiang, D Liu and H Ping. The first draft of the manuscript was written by P Xiang and H Ping. Critical revision of the manuscript for important intellectual content: H Ping, Y Liu and P Xiang. All authors commented on previous versions of the manuscript. All authors read and approved the final manuscript.

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Availability of data and materials

No datasets were generated or analysed during the current study.

Declarations

Ethics approval and consent to participate

Ethical approval was not needed because data were extracted from primary published studies in which informed consent was obtained by investigators.

Competing interests

The authors declare no competing interests.

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