Electronic Supplementary Material:

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Effects of Tadalafil Once Daily or On-Demand versus Placebo on Time to Recovery of Erectile Function in Patients after Bilateral Nerve-Sparing Radical Prostatectomy

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Supplementary Figure S1: Trial design

Screening

Double-blind treatment

Drug-free
washout

Open-label
tadalafil
5 mg OaD

Tadalafil 5mg OaD + Placebo PRN

Placebo OaD + Tadalafil 20mg PRN

Placebo OaD + Placebo PRN

≤ 6wk

2 mo

3 mo

4 mo

6 wk

3 mo

V1 V2 V3 V4 V5 V6 V7 V8 V9

Randomization

Baseline

Month 9

Month 10.5

Month 13.5

Primary endpoint

Abbreviations: nsRP, bilateral nerve-sparing radical prostatectomy; d, day; mo, month; OaD, once a day; PRN, “pro-re-nata”/on-demand; V, visit; wk, week.

Previously published in: Montorsi et al. 2014 [15].
Supplementary Figure S2 Patient Disposition

Screened
N = 583

Randomized
N = 423

ITT and Safety Population (N = 422)

Tadalafil OaD
N = 139

Completed double-blind treatment
N = 114

Completed drug-free washout
N = 105

Completed study
N = 98

Tadalafil PRN
N = 143

Completed double-blind treatment
N = 122

Completed drug-free washout
N = 117

Completed study
N = 112

Placebo
N = 141

Completed double-blind treatment
N = 115

Completed drug-free washout
N = 108

Completed study
N = 105

Most frequent reasons for discontinuation:

- Entry criteria not met: 23.1%
- Patient decision: 22.2%
- Adverse event: 19.4%
- Lack of efficacy: 15.7%
- Lost to follow-up: 13.0%

Abbreviations: ITT, intention-to-treat; OaD, once a day; PRN, “pro-re-nata”/on-demand; N, number of patients.

Previously published in: Montorsi et al. 2014 [15].
Supplementary Figure S3  Percent Patients Achieving IIEF-EF Score $\geq 22$

|                   | Month 9     | Month 10.5 | Month 13.5 |
|-------------------|-------------|------------|------------|
| **Odds Ratio [95% CI]; p-value** |             |            |            |
| OaD vs. Placebo   | 2.15 [1.16, 3.99]; **0.016** | 1.14 [0.63, 2.06]; **0.675** | 1.34 [0.79, 2.28]; 0.273 |
| PRN vs. Placebo   | 1.50 [0.79, 2.85]; **0.210** | 0.89 [0.48, 1.65]; **0.704** | 1.35 [0.80, 2.29]; 0.259 |

Abbreviations: CI, confidence interval; IIEF-EF, International Index of Erectile Function-Erectile Function domain score; n.s., not significant; OaD, once a day; PRN, “pro-re-nata”/on-demand.

Previously published in: Montorsi et al. 2014 [15].
**Supplementary Figure S4** Cox Proportional Hazard Model on Time to EF-Recovery (IIEF-EF≥22) during DBT

| Treatment effect: p = 0.038 | HR [95% CI] | p-value |
|---------------------------|-------------|---------|
| Tadalafil 5 mg OaD vs. placebo | 1.90 [1.16, 3.12] | 0.011 |
| Tadalafil 20 mg PRN vs. placebo | 1.47 [0.88, 2.47] | 0.140 |

**Age group effect: p = 0.223**

- < 61 vs. 61-68 years
  - 1.30 [0.85, 1.98] 0.223

Abbreviations: CI, confidence interval; DBT, double-blind treatment; EF, erectile function; HR, hazard ratio; IIEF-EF, International Index of Erectile Function-Erectile Function domain score; OaD, once a day; PRN, “pro-re-nata”/on-demand.