# Cost-Effectiveness of [<sup>177</sup>Lu]Lu-PSMA-617 in Patients With Progressive Metastatic Castration-Resistant Prostate Cancer

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# **KEY FINDINGS & CONCLUSIONS**

- <sup>177</sup>Lu-PSMA-617 demonstrated substantial incremental QALY gains versus both cabazitaxel and SOC, and incremental costs at list price that resulted in ICERs above UK willingness-to-pay thresholds.
- <sup>177</sup>Lu-PSMA-617 is an innovative treatment that can improve survival outcomes alongside more tolerable side effects, and it represents a clinically significant advancement in the management of mCRPC patients with high medical unmet need.

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# INTRODUCTION

- Prostate cancer (PC) is the second most frequently diagnosed cancer in men, leading to approximately 397,000 annual deaths worldwide.<sup>1</sup>
- Most PC patients develop metastatic castration-resistant

# **METHODS**

- A partitioned survival model was developed with 3 health states (progression free, progressed, and dead) to estimate costs and quality-adjusted life-years (QALYs) over a lifetime horizon (10 years) using a weekly cycle length.
- Efficacy and safety data for cabazitaxel were informed by several sources.
  - A network meta-analysis, comprising 6 randomised controlled trials,<sup>3-8</sup> was performed to assess the relative efficacy of rPFS and OS; the rPFS fixed-effects hazard ratio was used in the base-case analysis (Figure 1).

- PC (mCRPC) with bone metastasis leading to symptomatic skeletal events (SSEs) associated with worsening quality of life, morbidity, and survival.<sup>2</sup>
- Patients with mCRPC who progress following first-line docetaxel, are not medically suitable for docetaxel, or have previously been treated with docetaxel for non-metastatic PC have limited remaining treatment options.
- [<sup>177</sup>Lu]Lu-PSMA-617 (<sup>177</sup>Lu-PSMA-617), a prostate-specific membrane antigen (PSMA)-targeted radioligand therapy, is a new treatment option for patients with progressive PSMA-positive mCRPC.
- The phase 3 VISION trial showed that
   <sup>177</sup>Lu-PSMA-617 plus standard of care (SOC)
   significantly prolonged imaging-based radiographic
   progression-free survival (rPFS), overall survival (OS),
   time-to-first SSE, and time to worsening of health-related quality of life versus SOC.<sup>3</sup>
- The cost-effectiveness of <sup>177</sup>Lu-PSMA-617 compared with cabazitaxel and SOC was evaluated from the United Kingdom (UK) healthcare payer perspective.

- Health-state occupancy was governed by rPFS and OS curves.
- SSEs and treatment-emergent adverse events (AEs) were modelled, independent of health state.
- Patients (mean age = 70 years) received treatment with <sup>177</sup>Lu-PSMA-617 (up to 6 doses), cabazitaxel (up to 10 doses), or SOC; upon disease progression they could receive subsequent lines of anticancer therapy and supportive care.
- The VISION trial<sup>3</sup> was the primary source of efficacy and safety data for <sup>177</sup>Lu-PSMA-617 and SOC.
- Various standard parametric and flexible models were fit to the observed rPFS, OS, and time-to-first SSE data to extrapolate over the model time horizon.
- The base-case survival models were selected based on an assessment of visual fit and statistical fit to the trial data and a comparison of long-term extrapolations to external data and survival predictions elicited from clinical experts (Figure 1).
- Due to limitations with intertrial heterogeneity and the TheraP trial<sup>9</sup> not being powered to robustly investigate OS, an OS Kaplan-Meier curve for cabazitaxel derived from a real-world database analysis in the UK was used in the base-case analysis (Figure 1).
- Safety data were taken from the CARD trial<sup>4</sup>; due to the similarity in SSE incidence to the VISION trial, time-to-first SSE was modelled to be the same as
   <sup>177</sup>Lu-PSMA-617.
- Treatment-specific health-state utility values applied in the base case were derived from EQ-5D-5L data collected in the VISION trial<sup>3</sup> and taken from NICE TA391<sup>10</sup> for cabazitaxel (post-progression only).
- Costs were captured for a diagnostic test for PSMAscreening, drug acquisition and administration, AE and SSE management, healthcare visits and disease monitoring, and terminal care.
- Unit costs were sourced from standard UK sources with a 2020/2021 cost-year.

## RESULTS

• The base-case deterministic results for pairwise comparisons are presented in Table 1.

 Table 1. Deterministic Base-Case Results

- The base-case probabilistic results were similar; the cost-effectiveness planes are presented in Figure 2.
- One-way sensitivity and scenario analysis showed that the incremental cost-effectiveness ratios (ICERs) were most sensitive to methods used to model OS, the use of treatment-independent utilities, <sup>177</sup>Lu-PSMA-617 treatment exposure, and the application of concomitant treatments alongside <sup>177</sup>Lu-PSMA-617 (comparison with SOC only).

### Figure 1. Base-Case Survival Models

(A) Radiographic Progression-Free Survival



Treatment	Total cost (£)	Total QALYs	Incremental cost (£)	Incremental QALYs	ICER (£/QALY)
<sup>177</sup> Lu-PSMA-617	110,010	1.15	—	—	—
Cabazitaxel	47,366	0.62	62,644	0.52	119,852
SOC	29,326	0.71	80,684	0.43	185,572

### Figure 2. Cost-Effectiveness Plane



PSA = probabilistic sensitivity analysis.

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#### **Conflict of Interest**

VR, AK, and PJ are paid employees of Novartis. JB and HH are employees of RTI Health Solutions and were paid consultants to Novartis in connection with the development of this poster.

### **Author Contributions**

Study conceptualisation and methodology: JB, HH, and VR. Non-clinical data collection and analysis: JB and HH. Writing of the original draft: JB. Review and editing: HH, VR, AK, and PJ. All authors contributed to the poster and approved the submitted version.

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