Janssen Research & Development *

Clinical Protocol

Protocol Title

A Phase 3 Randomized, Double-blind, Placebo-controlled Study of Pasritamig (JNJ-78278343), a T-cell-redirecting Agent Targeting Human Kallikrein 2, + Best Supportive Care Versus Best Supportive Care for Metastatic Castration-resistant Prostate Cancer

KLK2-comPAS

Short Title
Pasritamig vs Placebo in late line Metastatic Castration-resistant Prostate Cancer (mCRPC)

Protocol 78278343PCR3001; Phase 3
Version: Original

JNJ-78278343 (pasritamig)

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Studies conducted at sites in the United States (US) will be conducted under US Food & Drug Administration Investigational New Drug (IND) regulations (21 CFR Part 312).
Studies conducted at sites in the European Economic Area (EEA) will be conducted under Regulation [EU] No 536/2014.
IND: 157066
EU TRIAL NUMBER: 2025-520927-26

Status: Approved
Date: 22 April 2025
Prepared by: Janssen Research & Development, LLC
EDMS number: EDMS-RIM-1382868, 2.0

GCP Compliance: This study will be conducted in compliance with Good Clinical Practice, and applicable regulatory requirements.

1. PROTOCOL SUMMARY

1.1. Synopsis

A Phase 3 Randomized, Double-blind, Placebo-controlled Study of Pasritamig (JNJ-78278343), a T-cell-redirecting Agent Targeting Human Kallikrein 2, + Best Supportive Care Versus Best Supportive Care for Metastatic Castration-resistant Prostate Cancer

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Pasritamig vs Placebo in late line Metastatic Castration-resistant Prostate Cancer (mCRPC)

Pasritamig (JNJ-78278343) is a humanized, IgG1-based, bispecific antibody designed to direct T cells (through binding to CD3 receptor complex) to hK2 (encoded by *KLK2* gene and hereafter referred to as KLK2) on target cells, leading to the activation of the T cells and T-cell-mediated lysis of KLK2-bearing cells.

OBJECTIVES AND ENDPOINTS

Primary objective: To determine if pasritamig + BSC compared to placebo + BSC is superior in OS.

Hypothesis

The hypothesis is that pasritamig + BSC will demonstrate improved OS, compared to placebo + BSC in participants with mCRPC.

OVERALL DESIGN

Key aspects of the trial design are summarised in this table.

Study Model:	Parallel group, 2 treatment groups	**Population Type:**	Adult patients
Control:	Placebo + BSC	**Population Diagnosis or Condition:**	mCRPC in the late-line setting
Study Treatment Assignment Method:	2:1 randomization, stratified by prior PSMA-targeted radioligand therapy, prior taxane, and ECOG performance status	**Population Age:**	≥18 years of age
Blinding	Double-blind	**Site Distribution:**	Multinational
Sub-studies	No		

Approximately 663 participants with mCRPC will be randomly assigned in this study in a 2:1 ratio to receive pasritamig IV + BSC or placebo IV + BSC. All participants must be receiving background ADT or had prior orchiectomy. All participants may receive BSC (defined as palliative external beam radiation, low dose steroids, pain medication, bone sparing agents, and needed palliative procedures) at the discretion of the physician.

The end of study/study completion is considered as the last scheduled study assessment for the last participant in the study.