

Contemporary Strategies for Clinical Chemoprevention of Localized Prostate Cancer

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Nagi B Kumar, PhD¹ 

Abstract

Prostate cancer (PCa) is the most common cancer among men in the United States and the second leading cause of cancer-related deaths. Metastatic castration-resistant PCa is still a fatal disease. On the other hand, between 2016 and 2020, about 70% of PCa cases were diagnosed at a localized stage. Evolving data demonstrates that men with low-grade cancers treated with definitive therapies may now be exposed to morbidities of overtreatment and poor quality of life, with little or no benefit in terms of cancer specific mortality. Active surveillance (AS) is thus the recommended management strategy for men with low-grade disease. Although this subgroup of men have reported anxiety during the AS period, they account to be highly motivated to make positive lifestyle changes to further reduce their risk of PCa progression, underscoring the urgent need to identify novel strategies for preventing progression of localized PCa to metastatic disease through pharmacologic means, an approach termed chemoprevention. Although several promising agents and approaches have been examined over the past 2 decades, currently, there are several limitations in the approach used to systematically examine agents for chemoprevention targeting men on AS. The goal of this review is to summarize the current agents and approaches evaluated, targeting men on AS, recognize the gaps, and identify a contemporary and comprehensive path forward. Results of these studies may inform the development of phase III clinical trials and ultimately provide a strategy for clinical chemoprevention in men on AS, for whom, currently, there are no options for reducing the risk of progression to metastatic disease.

Keywords

prostate cancer, clinical chemoprevention, active surveillance, biomarkers, systematic approach, % Ki-67, serum prostate specific antigen, gleason score, multiparametric MRI

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Introduction

Prostate Cancer

It is estimated that about 299 010 new cases of prostate cancer (PCa) will be diagnosed in the US in 2024, and 35 250 deaths will occur in men with PCa (ACS 2024).¹ Since the year 2014, the incidence rate of PCa has increased by 3% per year overall and by about 5% per year for advanced-stage PCa. PCa is the most common cancer among men in the US and the second leading cause of cancer-related deaths in the US. Some of the definitive risk factors associated to PCa includes age, genetic predisposition and race.² In 2019, 37% of PCa cases were

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