# 8th Global Summit on Plant Science

October 24-25, 2022

**WEBINAR** 

Ademola Emmanuel Adetunji, J Plant Physiol Pathol 2022, Volume 10

## <u>Capsaicin: A two-decade systematic review of global research output and recent advances against human cancer</u>

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apsaicin (8-methyl-N-vanillyl-6-nonenamide) is one of the most important natural products in the genus ✓ Capsicum. Due to its numerous biological effects, there has been extensive and increasing research interest in capsaicin, resulting in increased scientific publications in recent years. Therefore, an in-depth bibliometric analysis of published literature on capsaicin from 2001 to 2021 was performed to assess the global research status, thematic and emerging areas, and potential insights into future research activities. Furthermore, recent research advances of capsaicin and its combination therapy on human cancer as well as their potential mechanisms of action were described. In the last two decades, research outputs on capsaicin have increased by an estimated 18% per year and were dominated by research articles at 93% of the 3753 assessed literature. In addition, anticancer/pharmacokinetics, cytotoxicity, in vivo neurological and pain research studies were the keyword clusters generated and designated as thematic domains for capsaicin research. It was evident that the United States, China, and Japan accounted for about 42% of 3753 publications that met the inclusion criteria. Also, visibly dominant collaboration nodes and networks with most of the other identified countries were established. Assessment of the eligible literature revealed that the potential of capsaicin for mitigating cancer mainly entailed its chemo-preventive effects, which were often linked to its ability to exert multi-biological effects such as anti-mutagenic, antioxidant and anti-inflammatory activities. However, clinical studies were limited, which may be related to some of the inherent challenges associated with capsaicin in the limited clinical trials. This review presents a novel approach to visualizing information about capsaicin research and a comprehensive perspective on the therapeutic significance and applications of capsaicin in the treatment of human cancer.

#### Importance of Research

There is increasing interest in using capsaicin as a therapeutic alternative for different diseases (1,2) due to its pleiotropic pharmacological effects on various physiological systems, with an emphasis on pain as well as neuroscience, cardiovascular, respiratory, cancer, and urinary systems studies (3). With regards to pharmacokinetics, capsaicin has high oral bioavailability and skin absorption (4), making its topical application effective in various musculoskeletal or neuropathic pain conditions such as arthritis (5), shingles (6), vasomotor rhinitis (7), vasogenic facial pain (8). It is also used for treating urinary incontinence, chronic kidney disease-associated pruritus, and postoperative nausea and vomiting in acupoint therapy (9). Other beneficial bioactivities of capsaicin, including analgesic, anesthetic, anti-apoptotic, anti-inflammatory, anti-obesity, antioxidant, neuroprotective effects (10–12), enhanced energy metabolism (13), gastroprotective (14), and anticarcinogenic properties (15,16) have also been reported. However, capsaicin may also function as

Journal of Plant Physiology & Pathology

Volume 10

ISSN: 2329-955X

# 8<sup>th</sup> Global Summit on **Plant Science**

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a carcinogen or co-carcinogen (16–18). Understanding capsaicin research from global perspectives over an extended time is crucial. Although several studies have been published on capsaicin applications, bioactivities, and many other capsaicin-related topics (19–21), none of these studies has explored the scientometric approach to critically assess its progress and current direction in scientific research. Bibliometrics is a valuable tool for evaluating research trends within a subject area, thereby providing insight into extensively researched themes and identifying research needs to inform action (22). Here, an in-depth analysis of retrieved publications provided an overview of explored themes, research progression, as well as insights and future perspectives needed to enrich the knowledge domain on the compound.

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### **Biography**

Ademola has completed his PhD in Biology at the age of 35 years from the University of KwaZulu-Natal, South Africa. He is a postdoctoral research fellow at the Department of Molecular and Cell biology, University of Cape Town, South Africa where he co-supervises postgraduate students (Hons, MSc, and PhD) and oversees seed and cosmentic research projects. He has published 15 papers in reputed journals and has been serving as an invited reviewer for reputable journals.

Received: July 17, 2022; Accepted: July 19, 2022; Published: October 25, 2022

Journal of Plant Physiology & Pathology

Volume 10 ISSN: 2329-955X