

## **Overview**

Ivermectin, a well-known anti-parasitic drug, has garnered research interest for potential anticancer properties. While lab-based studies are promising, clinical validation is limited. This report provides a rigorous analysis of its mechanisms, supporting research, and current standing.

### **Mechanisms of Action (Preclinical)**

- Induces cancer cell death: Promotes apoptosis, autophagy, and pyroptosis across multiple cancer cell types.
- Inhibits proliferation pathways: Suppresses WNT/ $\beta$ -catenin, Akt/mTOR, and MAPK signalling to stop cell replication.
- Blocks metastasis: Inhibits PAK1 and other regulators of cytoskeletal rearrangement to prevent tumour spread.
- Overcomes chemotherapy resistance: Downregulates EGFR/ERK/Akt/NF- $\kappa$ B, restoring chemo sensitivity in resistant cells.
- Anti-angiogenesis: Reduces VEGF and other signals needed to form new blood vessels feeding tumour tissue.

### **Key Preclinical and In Vitro Studies**

- **Pancreatic Cancer**

Ivermectin in combination with gemcitabine caused G1 cell cycle arrest and apoptosis in human pancreatic cell lines.

Source: [https://aacrjournals.org/cancerres/article/82/12\\_Supplement/2320/701043/Abstract-2320-Ivermectin-suppresses-pancreatic](https://aacrjournals.org/cancerres/article/82/12_Supplement/2320/701043/Abstract-2320-Ivermectin-suppresses-pancreatic)

- **Esophageal Squamous Cell Carcinoma (ESCC)**

Mitochondrial dysfunction and ROS generation induced apoptosis in ESCC lines exposed to ivermectin.

Source: <https://bmccancer.biomedcentral.com/articles/10.1186/s12885-021-09021-x>

- **Colorectal Cancer**

Dose-dependent inhibition of proliferation and induced apoptosis in colorectal cancer cells.

Source: <https://www.frontiersin.org/articles/10.3389/fphar.2021.717529/full>

### **Clinical Trials and Human Data**

- **Triple-Negative Breast Cancer Trial (NCI-2022-02421)**

A Phase II trial testing ivermectin with pembrolizumab for metastatic TNBC. Status: Withdrawn.

Clinical Trials Registry: <https://www.cancer.gov/research/participate/clinical-trials-search/v?id=NCI-2022-02421>

## IVERMECTIN IN CANCER TREATMENT: SCHOLARLY EVALUATION

- Colorectal & Prostate Cancer Clinical Study (NCT05318469)

Investigates ivermectin monotherapy for tumour regression in a mixed cohort.

Registry: <https://clinicaltrials.gov/study/NCT05318469>

### Safety and Regulatory Status

- Approved as an anti-parasitic; not approved for cancer treatment by the FDA, EMA, TGA or other major regulatory bodies.
- Anticancer doses used in vitro often exceed safe human thresholds.
- Off-label use outside of trials may present toxicity risks.

### Expert Summary

1. Ivermectin shows credible anticancer potential in vitro and in vivo preclinical models.
2. Multiple mechanisms make it a candidate for multi-target therapies, especially when used synergistically.
3. However, there is no confirmed clinical efficacy from randomised controlled trials.
4. Its use should remain within ethically approved clinical trials only.
5. Caution is warranted when interpreting anecdotal or non-peer-reviewed claims online.

### Verified Reference Sources

[http://aacrjournals.org/cancerres/article/82/12\\_Supplement/2320/701043/Abstract-2320-Ivermectin-suppresses-pancreatic](http://aacrjournals.org/cancerres/article/82/12_Supplement/2320/701043/Abstract-2320-Ivermectin-suppresses-pancreatic)

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