

# Prediction of tumor growth at the microscopic level through mathematical modeling

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**Abstract.** All articles *must* contain an abstract. The abstract text should be formatted using 10 point Times or Times New Roman and indented 25 mm from the left margin. Set the pre-paragraph to 0 pounds and the post-break to 22.7 pounds. Starting on the same page as the abstract. The abstract should give readers concise information about the content of the article and indicate the main results obtained and conclusions drawn. The abstract is not part of the text and should be complete in itself; no table numbers, figure numbers, references or displayed mathematical expressions should be included. It should be suitable for direct inclusion in abstracting services and should not normally exceed 200 words in a single paragraph. Since contemporary information-retrieval systems rely heavily on the content of titles and abstracts to identify relevant articles in literature searches, great care should be taken in constructing both. To gain valuable insights into the growth of microscopic tumors, which is crucial for advancing cancer research, scientists employ mathematical modeling techniques. These models help researchers to comprehend how cells interact, proliferate, and organize spatially within tumors, ultimately aiding in developing better treatment strategies, including personalized medicine. A recent study introduced a novel approach using the Occam Plausibility Algorithm (OPAL) for modeling tumor growth. OPAL is a probabilistic framework adept at generating and refining hypotheses based on complex datasets. In this study, researchers integrated two existing models, the Proliferation Invasion Model (PIM) and the Mathematical Phase Field Model (MPFM), to create a comprehensive understanding of microscopic tumor growth dynamics. In this research, we have employed a literature review methodology to explore how researchers utilize three various mathematical modeling techniques, including the Occam Plausibility Algorithm (OPAL), to delve deeply into the growth dynamics of microscopic tumors. The conclusions drawn emphasize the significant potential of mathematical modeling in advancing scientists' understanding of microscopic tumor growth.

**Keywords:** Tumor Growth Prediction, Microscopic Tumor Modeling, Mathematical Simulation, Cancer Growth Patterns, Predictive Modeling In Oncology.

## 1. Introduction

Cancer is a highly intricate and diverse disease, and gaining insights into how tumors develop at the microscopic level is vital for creating effective treatments [1]. Recent advancements in mathematical modeling have opened up possibilities to simulate and forecast tumor growth dynamics. In the process of reviewing the literature, I discovered that three mathematical models, the Occam Plausibility

Algorithm (OPAL) [2], the Proliferation Invasion Model (PIM) [3], and the Mathematical Phase Field Model (MPFM) [4], are the most popular in the academic community for simulating and predicting the growth of microscopic tumors in the macro background. However, multiple studies in the literature have concluded that there is no one-size-fits-all model for predicting microscopic tumor growth. Nevertheless, by scrutinizing the strengths and weaknesses of these different models, it is possible to cultivate a more comprehensive understanding of tumor dynamics and identify the most suitable model for specific applications. This research delves into a comparison of three mathematical models used to predict microscopic tumor growth: the Occam Plausibility Algorithm (OPAL) [2], the Proliferation Invasion Model (PIM) [3], and the Mathematical Phase Field Model (MPFM) [4]. These models were harnessed to simulate tumor growth and evaluate their respective abilities to capture the complex facets of tumor dynamics, including cellular interactions, proliferation rates, migration angiogenesis, and spatial patterns, by literature review methodology. The study uncovered that each model possesses its unique strengths and limitations. OPAL serves as a robust tool for generating and refining hypotheses related to tumor growth, but it comes with the drawback of being computationally intensive. PIM offers a more detailed model capable of capturing the spatial and temporal dynamics of tumor growth; however, it may pose challenges in terms of parameterization. MPFM, while still in its developmental stages, holds potential as a more accurate and efficient approach to simulate tumor growth. The implications of this study for cancer research and clinical practice are significant. Researchers can devise more precise and efficient methods to simulate tumor growth and predict its dynamics by comprehending the strengths and limitations of various mathematical models. This knowledge can be harnessed to formulate personalized medicine and tailored treatment strategies with a higher likelihood of success. Furthermore, these findings can help bridge the gap between mathematical modeling and clinical decision-making. The development of user-friendly tools that facilitate clinicians' use of mathematical models can potentially enhance patient care and outcomes. This study contributes to the ongoing endeavors to unravel the complexities of cancer progression and enhance the effectiveness of therapies. Its findings hold promise for improving patient care and outcomes in the realm of cancer treatment.

## **2. Introduction to Mathematical Model**

Mathematical modeling is a formidable tool for comprehending the intricate nuances of microscopic tumor growth. Through the simulation of tumor development within a computer-based environment, researchers can unravel the complex interplay among cancer cells, the tumor microenvironment, and the immune system. Various mathematical models have been crafted to elucidate tumor growth, with differing focuses and approaches. Some delve into the realm of individual cells, while others adopt a broader perspective encompassing the entire tumor. Moreover, these models diverge in determinism, with some offering singular predictions for specific conditions [5], while others embrace stochasticity, accommodating multiple potential outcomes. The Occam Plausibility Algorithm (OPAL) stands tall among the esteemed mathematical models of tumor growth. Operating on Bayesian principles, OPAL employs probability theory to discern the most plausible explanation for a given dataset. It has found utility in investigating diverse tumor types, including breast, lung, and colorectal cancer. Another pivotal model is the Proliferation Invasion Model (PIM), which simulates the proliferation and dissemination of cancer cells within a three-dimensional environment. PIM accounts for crucial factors like cell division, migration, and apoptosis, and has been instrumental in studying the impact of various treatments on tumor growth. In the realm of innovative models, the Mathematical Phase Field Model (MPFM) emerges, positing that tumors are not uniform masses but comprise a mixture of cells and fluid. MPFM has been harnessed to scrutinize how the tumor microenvironment influences tumor growth. Mathematical models of tumor growth serve as invaluable assets in cancer research. They facilitate the assessment of novel treatments, enable predictions regarding disease progression, and contribute to formulating personalized therapeutic strategies. As mathematical modeling techniques continue to advance, their relevance in comprehending and combating cancer is poised to expand further. Beyond the models delineated above, an array of other mathematical models for tumor growth exists, each replete with its own merits and limitations. The selection of the most apt model for a specific study

hinges on the precise objectives at hand. However, all these models share the common virtue of shedding light on the intricate tapestry of tumor growth. With evolving technology and the accumulation of more data, mathematical modeling is destined to assume an even more pivotal role in cancer research. By furnishing deeper insights into tumor development, mathematical models have the potential to propel the development of more efficacious treatments and bolster the prospects of survival for individuals grappling with cancer.

### **3. Comparative Analysis of Mathematical Models for Microscopic Tumor Growth Prediction**

#### *3.1. The Occam Plausibility Algorithm model*

The Occam Plausibility Algorithm, known as OPAL, is a mathematical model that plays a crucial role in conducting comparative analyses of tumor growth models. Its Bayesian nature sets OPAL apart, relying on probability theory to discern the most likely explanation for a given dataset. This attribute renders it an invaluable instrument for identifying plausible interpretations of tumor growth patterns and for the selection, calibration, and validation of mathematical models. In a study conducted by Lima et al. in 2020 [2], OPAL was employed to compare three distinct tumor growth models: the Proliferation Invasion Model (PIM), the Mathematical Phase Field Model (MPFM), and the Individual-Based Model (IBM). OPAL's utility in this context lies in its ability to pinpoint the most probable explanation for the observed tumor growth patterns [6]. This entailed fine-tuning the parameters of the models to align with the actual tumor growth observations. The outcomes demonstrated that OPAL effectively calibrated the parameters of all three models, underscoring its value in ensuring the accuracy and reliability of mathematical models. Furthermore, OPAL was instrumental in validating these models, involving a comparison between their predictions and the real tumor growth patterns. The findings disclosed that both the PIM and MPFM accurately predicted the tumor growth patterns, whereas the IBM did not fare as well. This highlights that the PIM and MPFM stand out as more precise models for forecasting tumor growth compared to the IBM. This study's results underscore the significance of OPAL in facilitating comparative analyses of tumor growth models. OPAL's capabilities encompass identifying credible interpretations of tumor growth patterns, fine-tuning and validating mathematical models, and evaluating prediction accuracy [7]. In sum, OPAL emerges as a pivotal tool for advancing our comprehension of tumor growth and enhancing the development of more effective cancer treatments.

#### *3.2. Proliferation invasion Model*

OPAL, the Occam Plausibility Algorithm, is a mathematical model that is pivotal in the comparative analysis of tumor growth models. What distinguishes OPAL is its Bayesian foundation, leveraging probability theory to discern the most probable explanation for a given dataset[8]. This characteristic is a priceless utility in identifying credible interpretations of tumor growth patterns and in the selection, calibration, and validation of mathematical models. In a study conducted by Lima et al. in 2020 [2], OPAL was deployed to assess three distinct tumor growth models: the Proliferation Invasion Model (PIM), the Mathematical Phase Field Model (MPFM), and the Individual-Based Model (IBM). OPAL's prowess in this context lies in pinpointing the most likely explanation for the observed tumor growth patterns [6]. This involved fine-tuning model parameters to align with actual tumor growth observations. The results demonstrated OPAL's effectiveness in calibrating the parameters of all three models, reinforcing its role in ensuring the precision and trustworthiness of mathematical models. Furthermore, OPAL played a pivotal role in model validation, comparing their predictions against actual tumor growth patterns. The findings revealed that both the PIM and MPFM adeptly predicted tumor growth patterns, while IBM fell short in this regard. This underscores the superiority of the PIM and MPFM as more accurate models for forecasting tumor growth in comparison to IBM. The outcomes of this study underscore the significance of OPAL in facilitating comparative analyses of tumor growth models. OPAL's capabilities encompass not only the identification of credible interpretations of tumor growth patterns but also the refinement and validation of mathematical models, along with the evaluation of

prediction accuracy [7]. In summary, OPAL emerges as an invaluable tool for advancing our understanding of tumor growth and bolstering the development of more efficacious cancer treatments.

### *3.3. Mathematical phase field model*

The Mathematical Phase Field Model (MPFM) stands as a mathematical tool employed for forecasting tumor growth and the intricate process of new blood vessel formation, known as angiogenesis, at the microscopic level. In a study conducted by Xu et al. in 2016, the MPFM assumed a central role in comparing the predictive capabilities of various mathematical models concerning microscopic tumor growth and angiogenesis. The foundational concept behind MPFM lies in the recognition that tumors aren't uniform solid masses but rather comprise a mixture of cells and fluid [4]. This model considers critical factors such as cell proliferation rates, migration behavior, and the surrounding tumor microenvironment [9]. MPFM has found its application in studying diverse tumor types, from breast to lung and brain tumors. It has consistently demonstrated its ability to accurately forecast tumor growth and angiogenesis across various scenarios. Notably, one of the significant advantages of MPFM is its potential for personalized treatment. By considering the unique characteristics of an individual's tumor, MPFM can provide insights into how that specific tumor might respond to different treatment modalities. This valuable information can guide tailored treatment approaches, ultimately enhancing patient outcomes. MPFM emerges as a potent tool for unraveling and predicting the complex tumor growth and angiogenesis processes. Its utility extends to the realms of cancer diagnosis, treatment, and prevention, holding the promise of improving outcomes in these critical areas. The study results underscore that, in this particular case, MPFM outperformed other models in predicting tumor growth and angiogenesis, affirming its value in comprehending and forecasting these processes [10]. Leveraging MPFM within a comparative analysis of mathematical models is a valuable strategy for assessing the efficacy of different modeling approaches.

## **4. Reflection and Suggestion**

The study introduces three distinct models for predicting microscopic tumor growth: the Occam Plausibility Algorithm (OPAL), the Proliferation Invasion Model, and the Mathematical Phase Field Model. Each of these models exhibits promise in their predictive capabilities, yet they also come with their inherent limitations. OPAL, for instance, serves as a valuable starting point in the modeling process. However, it falls short when it comes to providing in-depth insights into the intricate details of tumor growth. On the other hand, the Proliferation Invasion Model excels in simulating the dynamic interplay of cellular dynamics and interactions but regrettably overlooks the crucial aspect of the tumor microenvironment. Similarly, the Mathematical Phase Field Model adeptly couples tumor growth with angiogenesis, yet it may not encompass the entirety of factors influencing this complex process. To enhance the accuracy and comprehensiveness of these models, future research should pivot toward their integration and the inclusion of more nuanced biological information. For instance, OPAL could be harnessed to generate hypotheses concerning tumor growth, which could be tested through the Proliferation Invasion Model or the Mathematical Phase Field Model. Additionally, the infusion of patient-specific data into these models could result in more precise predictions. Furthermore, it is imperative to translate the findings of mathematical modeling into practical clinical applications. This could be accomplished by crafting user-friendly software tools that empower healthcare professionals to assess individual patient data and make well-informed treatment decisions. By bridging the divide between mathematical modeling and real-world clinical scenarios, researchers hold the potential to markedly enhance patient outcomes. In adhering to these recommendations, researchers can propel the prediction of microscopic tumor growth through mathematical modeling, utilizing user-friendly software tools to integrate and analyze data from diverse sources. This journey may involve developing models encompassing the intricate interplay of genetic, environmental, and immunological factors that sway tumor growth dynamics. Additionally, conducting clinical trials to evaluate the effectiveness of treatment strategies rooted in mathematical modeling represents a crucial step forward. These efforts

collectively promise to deepen our understanding of microscopic tumor growth, potentially ushering in novel and more effective cancer treatments, and ultimately advancing patient outcomes.

## 5. Conclusion

The conclusions drawn from this study underscore the pivotal role of mathematical modeling, particularly through the utilization of models such as the Occam Plausibility Algorithm (OPAL), the Proliferation Invasion Model, and the Mathematical Phase Field Model, in advancing our comprehension of microscopic tumor growth. The study's findings unequivocally highlight the significant contribution of mathematical modeling in expanding our insights into the intricacies of microscopic tumor growth. Among these models, OPAL, along with the Proliferation Invasion Model and the Mathematical Phase Field Model, has emerged as a particularly valuable asset for capturing the multifaceted aspects of tumor dynamics, encompassing cellular interactions, spatial patterns, and the intricacies of angiogenesis. The synergy between these models promises to offer a more comprehensive understanding of tumor growth, potentially paving the way for novel clinical applications. Future research should be directed at addressing the limitations of these models. This could involve collecting more diverse and extensive clinical data to rigorously validate the models. Furthermore, the incorporation of longitudinal data to account for the evolution of tumors over time holds promise for enhancing the predictive accuracy of these models. Furthermore, there is a pressing need to bridge the chasm between mathematical modeling and clinical practice. Developing user-friendly software tools is a pivotal step in this direction. Such tools would facilitate the translation of research findings into practical insights for healthcare professionals, assisting them in crafting personalized treatment strategies for cancer patients. In sum, the conclusions drawn from these articles underscore the immense potential of mathematical modeling in advancing our understanding of microscopic tumor growth. However, they also emphasize the imperative of addressing limitations through diverse research, encompassing data collection, model development, and creating practical tools that can directly benefit clinical practice.

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