

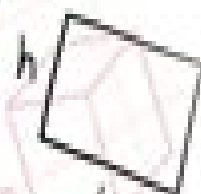


# IGCSE MATHS SOLUTION

## MAY / JUNE 2024

### QUESTION 2

### 0580/43/M/J/24



$$A = bh$$



$$V = s^3$$

M J 0580 43

**Thomas Griffiths**



**M J 0580 43:**

**Johannesburg** Transvaals,1960-07    Rapport från byggforskningen, Stockholm ,1977    New Publications of the U.S. Geological Survey ,1992    Metals Abstracts ,1996    New Publications of the U.S. Geological Survey Geological Survey (U.S.),1992    Military Law Reporter ,1982    **Dr. Dobb's Journal of Software Tools for the Professional Programmer** ,1977    **Contac** ,1994    **Johannesburg** ,1998    **Understanding convergent evasion mechanisms in cancer and chronic infection: Implications for immunotherapy** Matthias Theobald,Hansjörg Schild,2024-06-04 The complex interactions between the innate and adaptive immune systems function to recognize and clear pathogens or transformed cells but inefficient interactions between these two systems can result in harmful immunologic responses including chronic infections and the development of cancer Several hallmarks of dysfunctional adaptive immune responses often detected in tumors share specific features with ineffective immunity in chronic infections The members of the microenvironment actively participate in the process of tumorigenesis or chronification of infection by modulating innate and adaptive immune system interactions leading e g to insufficient T cell responses The best example is given by the acquisition of an exhausted state of cytotoxic CD8 T cells CTLs responding to chronic infections or tumors that are associated with elevated expression of inhibitory receptors and impaired cytokine response Targeting these major inhibitory pathways by immune checkpoint blockers represents a prime example of successful clinical translation of tumor specific immunotherapies Understanding the mechanisms behind mal adaptations of the immune system is crucial for achieving therapeutic benefits The establishment and co evolution of a dynamic microenvironment niche constituted by the recruitment of numerous cell types dampen immune responses and thus contribute to the development of neoplastic transformation as well as infection Although there are examples of successful immunotherapeutic approaches CAR T cells immune checkpoint inhibitors or mRNA vaccination a large percentage of patients with cancer or chronic infections still do not benefit from these therapies or develop severe immune related adverse events The reasons for these failures are not well understood A possible explanation might be that current immunotherapies target predominantly the effector arm of the immune system by trying to reactivate dysfunctional T cells but do not sufficiently address the influence of the innate immune system and the contributions of the tumor microenvironment TME niche The main problem we would like to address in this special issue is how inappropriate function of the innate immune system affects adaptive immunity and contributes to inefficient anti cancer immunity and chronification of infections The central goal is to provide a more precise understanding of the various common and novel immune evasion mechanisms in cancers and in chronic infections to obtain a detailed map of common and disease specific immune escape checkpoints To that aim we want to compile a wide array of interdisciplinary studies exploring a comparative and multi layered analysis of mechanisms responsible for inefficient immune responses including novel approaches i e multi omics or epigenetic signaling We would also like to combine studies from different fields including basic and clinical

immunology oncology and virology microbiology We welcome the submission of Original Research Review Mini Review Methods Case report and Perspective articles that cover but are not limited to the following topics Convergent mechanisms supporting immune escape in preclinical models tumors and chronic infections Convergent evasion mechanisms mediated by tumor infiltrating suppressive cells Treg MDSC macro phages soluble mediators signaling metabolism Convergent immune evasion mechanisms mediated by chronic infection viral or parasite Novel strategies to modulate the TME by direct or indirect targeting of immune suppressor cells Approaches to enhance persistence and resilience of anticancer T cells Combinatorial therapeutic strategies mRNA antibodies immune checkpoint blockers that target convergent immune evasion mechanisms Manuscripts consisting solely of bioinformatics or computational analysis of public genomic or transcriptomic databases which are not accompanied by robust and relevant validation clinical cohort or biological validation in vitro or in vivo are out of scope for this topic

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