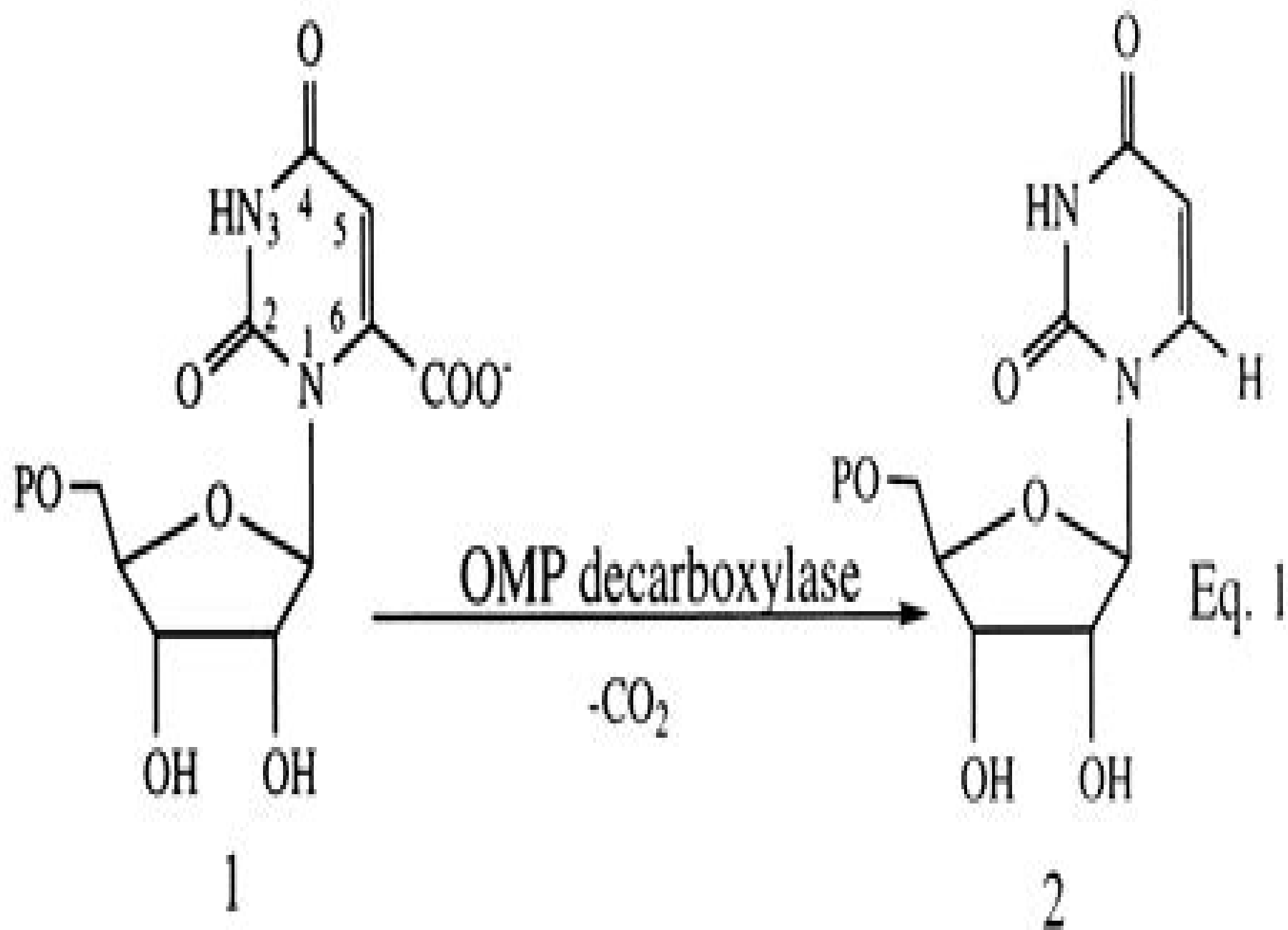




Orotidine Monophosphate Decarboxylase

Jeethiun K. Lee, Dean J. Tantillo



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Elucidating the Mechanism of Action of Orotidine Monophosphate Decarboxylase (ODCase) Using Isotope Labeling, Enzyme Kinetics and 6-CN-UMP Inhibitor Wickliffe O. Wepukhulu, 2006 Expression and Mechanistic Studies on Orotidine-5'-monophosphate Decarboxylase Kathryn J. Kitzmiller, 2004 **Mechanistic Study of Orotidine 5'-monophosphate Decarboxylase Using Uridine 5'-monophosphate Analogues** Freeman M. Wong, 2003

Investigations Into the Mechanism of Orotidine 5'-monophosphate Decarboxylase: Sources of Substrate Destabilization and Transition State Stabilization Vanessa A. Iiams, 2011 Orotidine 50 monophosphate decarboxylase OMPDC achieves a rarely paralleled rate acceleration yet the catalytic basis prompting this enhancement have yet to be fully elucidated To accomplish decarboxylation OMPDC must overcome the high energy barrier due to the localized anionic charge of the intermediate Mechanistic studies employing enzyme mutagenesis and product or intermediate analogues were used to investigate possible transition state stabilization by a carbene resonance structure Viability of the carbene structure depends upon a key hydrogen bond between O4 of the substrate and the amide backbone of a conserved serine or threonine Substitution of the conserved residue with Pro resulted in a k_{cat} of 1 M 1s 1 deletion of the FUMP O4 resulted in a product analogue that does not undergo H6 exchange or inhibit decarboxylation Hence indirect evidence reveals the O4 backbone interaction plays an important role for binding and catalysis OMPDC likely has honed multiple mechanisms to attain its remarkable catalysis The successful crystallizations of OMPDC a decade ago sparked hypotheses that structure and sequence conserved residues induced productive strain on the substrate enzyme complex Here we demonstrate a new source of stress a hydrophobic pocket adjacent to the OMP carboxylate that exhibits kinetic parameters characteristic of substrate destabilization Substitution of these residues with hydrophilic side chains by providing hydrogen bonding partners decreased k_{cat} by 10 to 10 40 3fold The same substitutions display very little change in the rate of product H6 exchange supporting that this hydrophobic pocket affects the substrate enzyme complex before the transition state We also provide evidence that hydrophilic residues can insert water molecules into the pocket with detrimental effects to catalysis **Class 4 Lyases I** Dietmar Schomburg, Ida Schomburg, 2001-10-23 The Springer Handbook of Enzymes provides concise data on some 5 000 enzymes sufficiently well characterized and here is the second updated edition Their application in analytical synthetic and biotechnology processes as well as in food industry and for medicinal treatments is added Data sheets are arranged in their EC Number sequence The new edition reflects considerable progress in enzymology the total material has more than doubled and the complete 2nd edition consists of 39 volumes plus Synonym Index Starting in 2009 all newly classified enzymes are

treated in Supplement Volumes **Enzyme Handbook** Dietmar Schomburg, Margit Salzmann, 2013-11-11 Recent progress in enzyme immobilisation enzyme production coenzyme regeneration and enzyme engineering has opened up fascinating new fields for the potential application of enzymes in a large range of different areas As more progress in research and application of enzymes has been made the more apparent has become the lack of an up to date overview of enzyme molecular properties The need for such a data bank was also expressed by the EC task force Biotechnology and Information Therefore we started the development of an enzyme data information system as part of protein design activities at GBF The present book Enzyme Handbook represents the printed version of this data bank In future it is also planned to make a computer searchable version available The enzymes in the Handbook are arranged according to the 1984 Enzyme Commission list of enzymes and later supplements Some 3000 different enzymes are covered Frequently very different enzymes are included under the same E C number Although we intended to give a representative overview on the molecular variability of each enzyme the Handbook is not a compendium The reader will have to go to the primary literature for more detailed information Naturally it is not possible to cover all numerous up to 40 000 literature references for each enzyme if data representation is to be concise as is intended *Antimalarial Agents* Graham L. Patrick, 2020-06-02 Antimalarial Agents Design and Mechanism of Action seeks to support medicinal chemists in their work toward antimalarial solutions providing practical guidance on past and current developments and highlighting promising leads for the future Malaria is a deadly disease which threatens half of the world's population Advances over several decades have seen vast improvements in the effectiveness of both preventative measures and treatments but the rapid adaptability of the disease means that the ongoing search for improved and novel antimalarial drugs is essential Beginning with a historical overview of malaria and antimalarial research this book goes on to describe the biological aspects of malaria highlighting the lifecycle of the parasite responsible for malaria the problem of resistance genetic mapping of the parasite's genome established drug targets and potential drug targets for the future This sets the scene for the following chapters which provide a detailed study of the medicinal chemistry of antimalarial agents with a focus on the design of antimalarial drugs Drawing on the knowledge of its experienced authors and coupling historic research with current findings to provide a full picture of both past and current milestones Antimalarial Agents Design and Mechanism of Action is a comprehensive yet accessible guide for all those involved in the design development and administration of antimalarial drugs including student academic researchers medicinal chemists malaria researchers and pharmaceutical scientists [Sulfur Amino Acids—Advances in Research and Application: 2013 Edition](#), 2013-06-21 Sulfur Amino Acids Advances in Research and Application 2013 Edition is a ScholarlyEditions book that delivers timely authoritative and comprehensive information about Cystathionine The editors have built Sulfur Amino Acids Advances in Research and Application 2013 Edition on the vast information databases of ScholarlyNews You can expect the information about Cystathionine in this book to be deeper than what you can access

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Foundations of Organic Chemistry David R. Dalton, 2011-08-04 This book differs from other organic chemistry textbooks in that it is not focused purely on the needs of students studying premed but rather for all students studying organic chemistry It directs the reader to question present assumptions rather than to accept what is told so the second chapter is largely devoted to spectroscopy rather than finding it much later on as with most current organic chemistry textbooks Additionally after an introduction to spectroscopy thermodynamics and kinetics the presentation of structural information of compounds and organic families advances from hydrocarbons to alcohols to aldehydes and ketones and finally to carboxylic acids

Enzymatic Reaction Mechanisms Perry A. Frey, Adrian D. Hegeman, 2007-01-27 Books dealing with the mechanisms of enzymatic reactions were written a generation ago They included volumes entitled Bioorganic Mechanisms I and II by T C Bruice and S J Benkovic published in 1965 the volume entitled Catalysis in Chemistry and Enzymology by W P Jencks in 1969 and the volume entitled Enzymatic Reaction Mechanisms by C T Walsh in 1979 The Walsh book was based on the course taught by W P Jencks and R H Abeles at Brandeis University in the 1960 s and 1970 s By the late 1970 s much more could be included about the structures of enzymes and the kinetics and mechanisms of enzymatic reactions themselves and less emphasis was placed on chemical models Walshs book was widely used in courses on enzymatic mechanisms for many years Much has happened in the field of mechanistic enzymology in the past 15 to 20 years Walshs book is both out of date and out of focus in todays world of

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Allosteric Regulatory Enzymes Thomas W. Traut, 2007-09-11 All enzymes are remarkable since they have the ability to increase the rate of a chemical reaction often by more than a billion fold Allosteric enzymes are even more amazing because they have the additional ability to change their rate in response to cellular activators or inhibitors This enables them to control the pathway in which they are the regulatory enzyme Since the effector molecules represent the current status of the cell for a given metabolic pathway this results in very responsive and balanced metabolic states and makes it possible for cells and organisms to be appropriately dynamic and responsive in a changing environment This book provides a logical introduction to the limits for enzyme function as dictated by the factors that are limits for life This book presents a complete description of all the mechanisms used for changing enzyme activity Eight enzymes are used as model systems after extensive study of their mechanisms Wherever possible the human form of the enzyme is used to illustrate the regulatory features Structural and Mechanistic Studies of Orotidine 5'-monophosphate Decarboxylase [microform] Ning Wu, 2002 **Class 4-6 Lyases, Isomerases, Ligases** Antje Chang, Dietmar Schomburg, Ida Schomburg, 2010-01-08 The Springer Handbook of Enzymes provides concise data on some 5 000 enzymes sufficiently well characterized and here is the second updated edition Their

application in analytical synthetic and biotechnology processes as well as in food industry and for medicinal treatments is added Data sheets are arranged in their EC Number sequence The new edition reflects considerable progress in enzymology the total material has more than doubled and the complete 2nd edition consists of 39 volumes plus Synonym Index Starting in 2009 all newly classified enzymes are treated in Supplement Volumes Cofactor Biosynthesis: A Mechanistic Perspective,2001-01-02 The longest running serial published by Academic Press continues its well respected run with Volume 61 a special volume in which a guest editor has come on board and has assembled some well known contributors who are international authorities in the field Together they tackle some of the latest topics in the field such as riboflavin and folate biosynthesis biotin and lipoic acid biosynthesis nicotinamide adenine dinucleotide biosynthesis biosynthesis of vitamin B6 and structurally related derivatives pantothenic acid and coenzyme A biosynthesis mechanistic biosynthesis of protein derived redox cofactors ascorbic acid biosynthesis biosynthesis of menaquinone and ubiquinone Vitamin B12 biosynthesis biosynthesis of the methanogenic cofactors and thiamin biosynthesis *Investigating Structural Dynamics of Orotidine 5'-monophosphate Decarboxylase* Andrew H. Shabila,2009

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