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STUDY OF PHOSPHATE AND PHOSPHONATES AS AN INTRIGUING SUBSTANCES FROM A WATER TREATMENT PERSPECTIVE

¹Author: Ashakiranmayi Siddavarapu, research scholar at the department of Chemistry at Sri Satya Sai University of Technology & Medical Sciences, Sehore-MP.

²Author: Dr. Neelu Jain, Professor at the department of Chemistry at Sri Satya Sai University of Technology & Medical Sciences, Sehore-MP.

Abstract:

As phosphate isosteres, a number of nucleoside phosphonates have been synthesised and studied. Examples of this type of compound include deoxy nucleoside phosphonates, which either remove or replace the nucleoside phosphorus position's oxygen with a methylene moiety. As the most promising isoster of the naturally occurring phosphonooxymethyl group (P–O–C), phosphonomethoxy (P–C–O) occurs when the 5'-oxygen and 5'-carbon are exchanged. As a result of the fact that it's isopolar and isosteric to phosphoric acid, its success can be explained. As a substrate for various kinases, nucleoside phosphonates can be phosphorylated by enzymes, which results in the formation of diphospho phosphonate analogues, is another key property of nucleoside phosphonates. In particular, this is crucial for antiviral and anticancer medicines, because in these situations, the pharmacologically active species is a nucleoside triphosphate analogue, which is not present in the natural product.

Keywords: phosphate, diphospho, wastewater, treatment, herpesviruses, etc.

1. INTRODUCTION

Phosphate and phosphonates are both unequivocally adsorbed onto mineral surfaces and their expulsion amid wastewater treatment is fundamentally because of adsorptive procedures. We have directed investigations to consider the common impact of phosphate and six distinctive phosphonates on each other in supported medium at pH 7.2. We have utilized phosphonates having one to five phosphonic corrosive gatherings (HMP, IDMP, HEDP, NTMP, EDTMP and DTPMP). The nearness of phosphonates stifled the adsorption of phosphate. The monophosphonate HMP had the littlest and the polyphosphonates the biggest impact on phosphate adsorption. The nearness of phosphate brought down

phosphonate adsorption. The opposition in the multicomponent framework can sensibly well be anticipated utilizing a surface complexation show produced for single segment systems. Phosphate and phosphonates are intriguing substances from a water treatment perspective. Phosphate released into surface waters can empower plant development, bringing about an eutrophication of waterways and lakes. A productive evacuation of phosphate amid wastewater treatment is along these lines an essential factor in saving the water nature of eutrophic lakes. Phosphonates are complexing specialists containing at least one C–PO(OH)₂ gatherings. Phosphonates have three primary properties: they are successful

chelating specialists for two- and tri-valent metal particles they hinder precious stone development and scale arrangement and they are very steady under brutal synthetic conditions.

2. ANTI-HERPESVIRAL NUCLEOSIDE PHOSPHONATES

All around the world, herpesviruses (of the Herpesviridae family) cause latent and recurring infections in many organs and tissues. Herpes simplex virus types 1 and 2 (HSV-1 and HSV-2), varicella-zoster virus (VZV), human cytomegalovirus (HCMV), human herpesvirus types 6A, 6B, and 7 (HHV-6A, HHV-6B, and HHV7), and the oncogenic Epstein-Barr virus (EBV) are among the nine types of herpesviruses known to routinely use humans as their primary host (KSHV). There have been reports of humans contracting a herpesvirus carried by macaques known as B virus. Immunocompromised individuals are at a disproportionately high risk of morbidity and death from human herpesvirus diseases. After discovering acyclic phosphonates, which were used to treat herpes infections, researchers in the 1980s identified nucleoside analogues that had an acyclic pseudo-sugar moiety (such as acyclovir, penciclovir, ganciclovir, and penciclovir). A major advance in herpes virus therapy was achieved (ANPs). Purine and pyrimidine nucleoside side chains can be connected to the phosphonate moiety of acyclic nucleosides (ACNs) through an irreversible P-C bond. However, ANPs do not require the presence of

a virally-encoded kinase to activate them. According to their chemical structure, ANPs are divided into subclasses, each of which has a specific antiviral activity range. HPMPs, for example, have a 2'-hydroxymethyl group, which allowed the development of broad-spectrum antiviral medicines that were both effective and selective against a wide variety of herpesviruses and other double-stranded DNA viruses (pox-, polyoma-, papilloma-, and adenoviruses). There is a preference for the (S)-enantiomers over the (R)-enantiomers in the C2' stereochemistry of HPMPs, which has a major effect on their antiviral activity.

- **Tautomeric equilibria**

Also of note, mono- and diesters of phosphonic acids occur in solution in an equilibrium combination of two tautomeric forms: the five-coordinate phosphonate form (5, 4), and the three-coordinate phosphite form (3, 3), respectively (Figure 1.5). Despite the fact that both compounds contain phosphorus in the +3 oxidation state, their chemical reactivity is fundamentally different. Due to the presence of an unpaired electron pair on the phosphorus atom, it is possible to react fast with diverse electrophiles by reacting with trivalent phosphorus (III) derivatives. These compounds are less sensitive to electrophiles because they lack a lone electron pair on the phosphorus centre. As a result, they are more stable and resistant to spontaneous oxidation than other phosphonates



Figure 1: Indices λ and σ stand for the valency and the coordination number of the phosphorus atom, respectively.

- **High electrophilicity at the phosphorus center**

When it comes to reactivity, H-phosphonate diesters are known for their high sensitivity to hydrolysis in acidic situations. Electrically neutral and base labile phosphate triesters hydrolyze at 105 times faster rates with this approach. In addition, activated H-phosphonate monoesters are very reactive, as seen by their fast condensation with alcohols when exposed to light. This increased reactivity of H-phosphonate derivatives is usually attributed to the presence of P–

H bonds in the compounds' structure. This theory, however, has been contested. The electronic effect of the P–H bond and the ability of these compounds to form tervalent species remain viable explanations, given that it is highly unlikely that low steric hindrance (resulting from the presence of the P–H bond) can result in such rapid acceleration of the rate of reaction in acyclic compounds. As a result, tetracoordinated H-phosphonic-carboxylic mixed anhydrides are much more reactive than tervalent bisacylphosphites

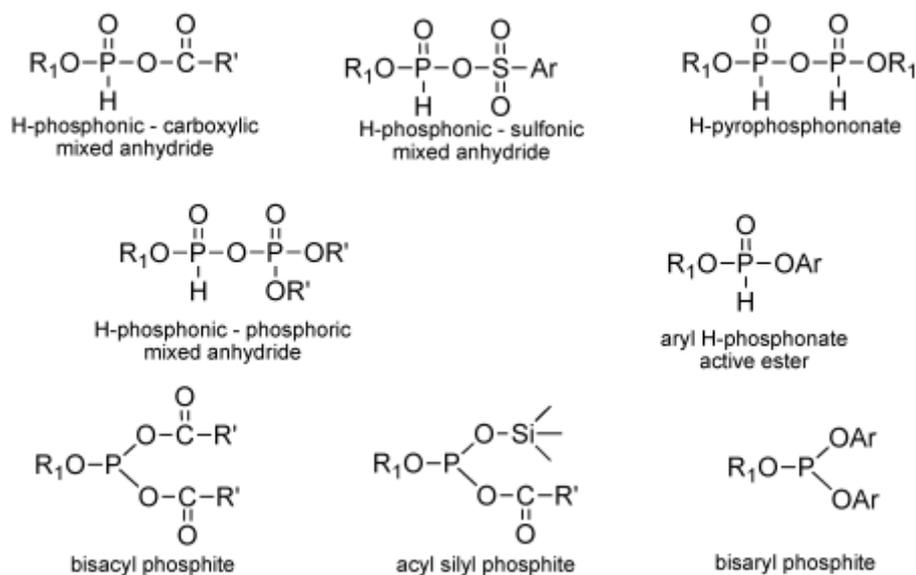


Figure 2: R_1 = a nucleoside moiety or carbohydrate, lipid, alkyl, aryl, etc. R' = alkyl or aryl.

It's not understood what causes this electrical effect; nonetheless, it's most likely related to phosphorus's chemical bonding system, which is dominated by back-donation from the phosphorus's substitutes. Due to the difference in electronegativity between oxygen (2.1) and phosphorus (2.1), the P–O bonds in phosphate triesters $[O=P(OR)_3]$ are directed towards oxygen (3.5). This is owing to the fact that phosphorus and oxygen have different electronegativity (3.5).

3. PHOSPHONATE BIOCHEMISTRY AND THE DEVELOPMENT OF PHOSPHONATES RESULTA AGRICULTURAL CHEMICALS

When political turbulence in Europe in the 1930s and the Pacific war in the 1940s threatened the supply of rock phosphate to farmers in the United States and Europe, the United States and Europe founded the Rock Phosphate Association. As early as the 1990s, German agronomists and their American counterparts at the U.S. Department of Agriculture

began looking for alternatives to phosphate fertilisers. It was expected that salts of phosphonic acid would release phosphorus more slowly than other compounds, and would therefore be more cost-effective in terms of delivering phosphorus to the plant. However, it was found that they were less effective than standard sources of phosphorus at stimulating plant growth. So, for the next three decades, phosphonates were exempted from agricultural duties. The fact that plants that had been treated with phosphonate in the laboratory showed delayed growth response is notable. Meanwhile, phosphonic acid salt and esters were also researched by biochemists in the 1950s, when they were conducting an exceedingly thorough investigation into the mechanism of oxidative-phosphorylation at the time of their discovery. Because these compounds were physiologically inactive, they may be used in cell-free conditions where phosphates would interfere with their function as a buffer, according to the research team. C-P phosphonates, on the other hand, have been found to possess

significant biological activity. Phosphonoacetic acid (foscarnet) and phosphonoformic acid (foscarnet) are antiviral medications, whereas diphosphates are used to manage bone demineralization. In addition to insecticides, several fungicides are also phosphonates, as previously mentioned. Growing control agents and herbicides use phosphonates. Phosphonic acid salts or esters in living creatures have yet to be detected in a scientifically verified manner. Natural phosphonates with C-P bonds, on the other hand, come in a wide range. All sorts of organisms have been used to separate them including protozoa, mollusks, and colenterates. According to some accounts, oomycetes also contain phosphonates. When it comes to C-P bonds, 2-amino ethyl phosphonate is the most common, as it is found in phosphonolipids as analogue of phosphoethanolamine. As a component of phosphonolipids, it is commonly encountered. C-P links in phosphonolipids are more resistant to chemical hydrolysis than C-O-P bonds, and phospholipase D does not break down phosphonolipid C-P bonds. Natural phosphonolipids have eluded explanation until recently, and their seemingly random distribution throughout the different kingdoms and phyla of the living world has been puzzling. Organisms with these compounds in their membranes and phospholipases as part of their invasion or swallowing of their host have been postulated. prey may have a competitive edge over other species. The presence of these enzymes should cause phosphonolipid-containing membranes to become more stable, as previously stated. However, as of now, there is little experimental data to back up this assertion, as previously stated.

Proteins, glycans, and some antibiotics generated by bacteria and fungus are examples of naturally occurring molecules that have been found to have a C-P link. Bialophos is unique in that it has a C-P-C bond, which makes it a rare compound. Several routes for the synthesis and degradation of 2-aminoethyl phosphonate have now been identified. Furthermore, the biosynthetic route for the compound bialophos has been identified. The Furthermore, compounds containing C-P bonds have attracted more attention, as many of these compounds are now used as insecticides, weedkillers and detergent ingredients as well as anti-inflammatories, to name just a few. Only bacteria have been identified to possess C-P bond cleaving enzymes. Genes for alkylphosphonate absorption and C-P lyase activity in *Escherichia coli* have only recently been identified.

4. INDUCED AS WELL AS NATURAL VARIABILITY IN PHOSPHONATE SENSITIVITY

Because of the significant differences found in sensitivity of *Phytophthora* species to phosphonates and alkyl phosphonates in axenic culture across different medium and development phases, it has been challenging to determine the sensitivity of these species to phosphonates and alkyl phosphonates. A study conducted on certain, but not all, species of bacteria revealed that the amount of phosphate present in their culture medium *Tvas* was a deciding factor in their sensitivity to phosphonate. Agar (CMA), which contains 0.38 msi of phosphate, has been used in the majority of following investigations of growth inhibition *in vitro*. Modified Ribeiro's format, which contains 0.38 msi of phosphate, has also been used in the majority of subsequent research (0.084 mM). *Phytophthora capsici* Leonian mycelial growth is reduced by phosphate deprivation alone at the phosphate concentration in modified Ribeiro's medium in the absence of phosphonate when the phosphate concentration in the medium is the same as the concentration in the media. On these medium, the inhibition of mycelial development is enhanced, allowing for more accurate ED measurements. Agar preparations with phosphate concentrations ranging from 1 to 1 million micrograms per litre might change the apparent susceptibility of a bacterial strain to phosphonate when they are used in solid media production. Furthermore, measures of inhibition based only on radial growth might be deceiving in their accuracy. There are phosphate and phosphonate combinations available that significantly lower the density of mycelia of *Phytophthora palmicora* and *P. infestans* on agar medium, while having no effect on radial growth rates on agar media (Figure 1). (Griffith, Davis, Snow and Grant, unpublished). It comes as no surprise, then, that *Phytophthora* species' sensitivity to phosphonate is inconsistent when measured in phosphate-limiting media, as opposed to plant tissues, where phosphate concentrations are typically 5-20 mxi, or that their sensitivity to phosphonate *in vivo* is inconsistent with their sensitivity when measured *in vitro*.

5. PHOSPHONATESTRANSLOCATION

The phosphonates are genuinely systemic fungicides, as opposed to other types of fungicides. The phloem and xylem of plants, unlike other systemic fungicides, are translocated by these fungicides. Phosphonate is rapidly absorbed and translocated in

the xylem immediately upon administration of the solution. Once in the phloem, it travels into and is translocated, and its distribution in the plant is subject to the typical source-sink connections that exist in the plant. In the field of disease control, the importance of source-sink interactions controlling phosphonate translocation for optimal disease control, particularly in perennial crops, is only now beginning to be recognised. It became feasible for the first time to use foliar sprays as both a preventive and curative therapy for root infections. Furthermore, it was discovered that, due to their high water solubility, selectivity, and genuine systemic mobility, phosphonates may be injected directly into tree trunks to manage root infections, which was a significant discovery. Avocado trees were the original target of this technique, which was designed to guard and heal them from Infections caused by *Phytophthora cinnamomi*. In recent years, it has been applied to a variety of tree crops. *Phytophthora cinnamomi* root rot in avocados is best controlled by trunk injections of phosphonates in late summer, when the leaf and fruit flush has stopped and the roots have become the plant's principal metabolic sinks, according to many ingenious studies. Same thing with cocoa, where early-season pesticide injections during a period of high disease pressure allow beans, which serve as key metabolic sinks during that time period, to accumulate phosphonate, while infection develops in the pod case (Guest, unpublished results). There should be more research done to discover if earlier injections of black pod control boost black pod control when the pod case is the principal metabolic sink for the pod. A thorough understanding of the host physiology and the source-sink interactions at various stages of plant growth is required to improve disease management in perennial crops by employing phosphonates.

When an organism is fed phosphate, the rate at which phosphonate is absorbed and, therefore, the effectiveness of the chemical in issue appears to be affected. Previous theories that oomycetes were more susceptible to the toxic effects of phosphonates were incorrect, as the ability of oomycetes to metabolise phosphonates differs from other species. Oomycetes have at least two phosphate uptake mechanisms, which is common to many other microorganisms and plants. As an example, one sort of system has a low affinity for phosphate and is active throughout a broad range of phosphate concentrations. According to earlier information, the other system is a high-affinity system that can only be generated under phosphate-limiting conditions. All dosages of

phosphonate and phosphonate are antagonistic in their fight for binding sites on transporter proteins. Phosphate absorption can be inhibited by phosphonate concentrations as low as μM concentrations, but not by mM concentrations of phosphonate (and vice versa). Also, in order for plants to absorb phosphates, they must use a phosphate carrier system. Based on research employing leaf discs it has been postulated that phosphate inhibition does not alter this process.

6. CONCLUSION

In spite of the fact that there were problems in the formation of symmetric salts, studies into the chiral trisphosphonate were initiated. While the production of various phosphites was relatively straightforward, the preservation of chlorophosphites was more problematic due to their highly reactive nature. Experiments with a dichloro- or trichlorophosphite to produce the trisphosphonate were unsuccessful due to the low yields of the product. The creation of the parent asymmetric bisphosphonate, on the other hand, was easier to regulate, permitting the use of a chlorophosphite that was readily accessible on the market.

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