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Microwave-assisted nucleophilic degradation of organophosphorus pesticides in propylene carbonate†

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Propylene carbonate is becoming a suitable green alternative to volatile organic solvents in the study of chemical reactions. In this study, an efficient method for nucleophilic degradation of five organophosphorus pesticides, fenitrothion, malathion, diazinon, parathion, and paraoxon, using propylene carbonate as a solvent is proposed. The effect of changing the nature of the nucleophile and the influence of microwave (MW) heating were investigated. A screening of temperatures (50 °C–120 °C) was performed under microwave heating. The pesticide degradation was followed by ³¹P NMR, and the extent of conversion (%) was calculated by the integration of phosphorus signals. Keeping in mind that recently it has been reported that some ionic liquids play a nucleophilic role, in this work we report for the first time the degradation of organophosphorus pesticides by using an amino acid-based ionic liquid such as Bmim [Ala] as a nucleophile and a bio-based solvent (propylene carbonate) as a reaction medium in combination with microwave heating.

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1. Introduction

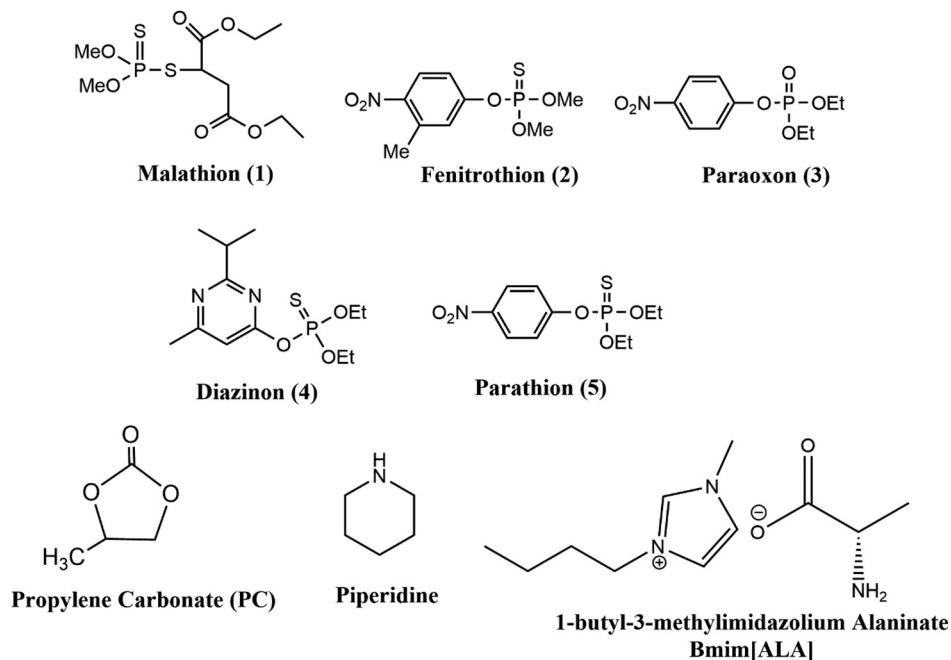
In the last few decades, the requirements for sustainable and safe processes have been gaining much attention, not only in academics and the chemical industry but the whole society. This trend towards what has become known as sustainable technologies can be defined as green chemistry which is driven by twelve principles described by Anastas.¹ Effort has been made to meet the demands of green chemistry principles mainly in the replacement of volatile organic solvents (VOS) by suitable alternatives such as biodegradables and non-volatile green solvents.^{2–4} In this context, very recently, bio-based solvents, which are derived from natural resources, such as vegetable and animal raw materials, have emerged as a new class of green solvents.^{5–8} They have been used as reaction media in some organic processes and have shown to be effective without compromising the safety and the environmental acceptability of the process.^{9–11} It is interesting to note that propylene carbonate (PC) is nowadays one of the greenest

solvents^{12,13} and it has been used as reaction media in many useful organic transformations.^{14–20} PC is known as a carbon dioxide neutral solvent and can be obtained from propylene oxide and carbon dioxide²¹ and also from a microwave-assisted process intensification of lipase-catalyzed transesterification of 1,2 propanediol with dimethyl carbonate.²²

On the other hand, organophosphorus pesticides (OPP), which are one of the most used pesticides worldwide, have increased the danger of their bioaccumulation in soils and groundwater.^{23–25} As a result, the degradation of these kinds of pesticides is an important issue to study. In this context, our group has been investigating the degradation of OPP by nucleophilic attack in several kinds of solvents such as aqueous solutions,²⁶ volatile organic solvents (VOS), ionic liquids (ILs) and bio-based solvents.^{27–29} For example, the nucleophilic degradation of fenitrothion with piperidine in PC at room temperature was very efficient ($t_{1/2}$ = 9 min), while the degradation of fenitrothion in Bmim[NTF₂] was quite similar ($t_{1/2}$ = 13.2 min) and much more slow in ethyl lactate ($t_{1/2}$ = 210 min).

It is well known that the efficiency of nucleophilic degradation of organophosphorus pesticides depends strongly on the nucleophile, the nature of the phosphoryl center of the pesticide, and the type of solvent used in the reaction. In this context, we have studied the nucleophilic degradation of paraoxon in a series of ILs containing 1-butyl-3-methylimidazolium (Bmim)⁺ as a cation and different amino acids (AA) as anions (Bmim[AA]).²⁹ It was observed that the reaction was completed at room temperature in the absence of an external nucleophile

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Therefore, in this study we will investigate the influence of microwave heating on the degradation of OPP with Bmim[Ala] and piperidine as nucleophiles using PC as the solvent. The pesticides malathion (1), fenitrothion (2), paraoxon (3), diazinon (4) and parathion (5) were chosen based on their electrophilic center phosphate (P=O) or phosphorothioate (P=S) with different leaving groups (Scheme 1).

2. Materials and methods

2.1. Materials

Propylene carbonate (PC), piperidine and organophosphorus pesticides were purchased from Aldrich. Piperidine was distilled before use. Bmim[Ala] was prepared as described.²⁹ PC was stored over molecular sieves.

2.2. Kinetic measurements

Kinetic measurements were performed by ^{31}P NMR using a 400 MHz spectrometer, following the disappearance of the pes-

ticide signal. The kinetic experiments were carried out in PC as the solvent at 25 °C, and Bmim[Ala] and piperidine as nucleophiles. The spectra were recorded at different reaction times and pseudo-first-order rate coefficients (k_{obsd}) were obtained by integration of the NMR signals for pesticides and plotting \log (integration) *vs.* time. Each measurement was made in triplicate, and the k_{obsd} values reported in Table S1, in the ESI† correspond to the average of the three measurements. In a typical experiment, a NMR tube containing 500 μL of the PC was thermostatted at 25 °C for 10 min, and then 10 μL of pesticide (0.5 M in ACN) and 20 μL of neat piperidine or 40 μL of Bmim[Ala] were added. A capillary tube of deuterated water was used as a reference solvent for NMR experiments.

2.3. Microwave-assisted degradation

Microwave-assisted reactions were carried out in an Anton Paar Monowave 300 Microwave Synthesis Reactor (Anton Paar GmbH, Graz, Austria) in 10 mL sealed vials by adding 500 μL of solvent, 10 μL of pesticide (0.5 M in MeCN) and 20 μL of neat piperidine or 40 μL of Bmim[Ala]. Reaction temperatures in the range of 50–120 °C for 10–60 minutes at 500 rpm were selected. To evaluate the percentage of conversion (%) of the pesticide, the reaction mixture was analyzed by ^{31}P NMR.

2.4. Gas chromatography–mass spectrometry (GC/MS)

GC/MS analysis was performed using a gas chromatograph fitted with a split–splitless injector and an Elite-5MS column (30 m \times 0.25 mm i.d., 0.25 μm d.f.). Helium was used as a carrier gas at a flow rate of 1.0 mL min^{-1} . The injection port was maintained at 25.0 °C, and the split ratio was 19 : 1. The oven temperature was programmed from 70 °C for 2 min, then ramp 1: 5 °C min^{-1} to 140 °C hold for 12 min and ramp 2: 10 °C min^{-1} to 240 °C hold for 12 min. The ionization mode was electron impact ionization and the scanning range was from 50 amu to 400 amu. Mass spectra were obtained at 0.35 s intervals.

2.5. Electrospray ionization mass spectrometry (ESI-MS)

The identification of the reaction products was performed using an ABSciex Triple Quad 4500 (UHPLC-MS/MS) mass spectrometer equipped with a Turbo Ion Spray (AB Sciex) ion source. A microsyringe pump delivered the reaction mixture of the pesticide with piperidine in propylene carbonate at infinite time dissolved in 10% (v/v) acetonitrile into the ESI source at a flow rate of 10 μL min^{-1} . ESI and QQ (linear trap) mass spectrometry were performed in the negative-ion mode for detecting organophosphate compounds by using the multiple reaction monitoring (MRM) scan type. Main conditions: curtain gas nitrogen flow 10 mL min^{-1} ; ion spray voltage –4500 eV; declustering potential –60 eV; entrance potential –10 eV; collision cell exit potential –12 eV; source temperature was set at 300 °C and source gases GS1 and GS2 were set to 12 and 0, respectively. All data were acquired using Analyst 1.6.2 (AB Sciex).

3. Results and discussion

It was previously described that Bmim[AA]-type ionic liquids have a dual function, as nucleophiles and solvents, in the degradation of pesticides.²⁹ In this study, we investigated the degradation of pesticides 1–5 (Scheme 1) using propylene carbonate (PC) as the solvent and Bmim[Ala] as the nucleophile (0.4 M) under pseudo-first order conditions. The reactions were followed by ^{31}P NMR experiments, and the degradation of the pesticides was calculated by the integration of the phosphorus signals and expressed as percentage (%) of extent of conversion.

First, the reaction of pesticides with Bmim[Ala] in PC was explored at room temperature, and it was found that fenitrothion, malathion and paraoxon were fully degraded, but not diazinon and parathion (Fig. S1–S3 in the ESI†). For comparison purposes, we then tested the reactivity of the five pesticides with a known nucleophile such as piperidine in PC at room temperature. The results showed that fenitrothion (2) and malathion (1) were quickly degraded (Fig. S4 and S5 in the ESI†), but paraoxon, parathion, and diazinon did not react even after 24 hours of reaction. From the plots of \log % degradation *vs.* time, we first calculated the pseudo-first order constants (k_{obs}) and then half-life ($t_{1/2}$) values which are shown in Table 1.

The best rate of degradation was found for malathion with both nucleophiles, which could be due to the fact that aliphatic pesticides are degraded faster than aromatic pesticides together with a loss of stability in OPP with –OMe groups.³⁶

We can observe that both nucleophiles can degrade fenitrothion, but piperidine is much more efficient. Noteworthy, malathion degradation with both nucleophiles shows similar half-life ($t_{1/2}$) values, which is very interesting due to the fact that we can use Bmim[Ala] without compromising the rate of the reaction. But the most promising result is that observed for paraoxon degradation when Bmim[Ala] is the nucleophile, since this nucleophile achieves a total degradation of this pesticide at room temperature ($t_{1/2}$ = 1220 min, Fig. S3 in the ESI†). These results confirm that Bmim[Ala] can be used as a reagent for the degradation of malathion (1) and paraoxon (2) at room temperature using a green and sustainable solvent such as PC.

Table 1 Half-lives ($t_{1/2}$) for the degradation of pesticides 1–5 in PC at room temperature

	Half-life ($t_{1/2}$) (min)	
	Bmim[Ala]	Piperidine
Malathion (1)	10.7 \pm 0.60	9.9 \pm 0.6
Fenitrothion (2)	425.0 \pm 11.4	23.1 \pm 0.6
Paraoxon (3)	1203 \pm 96.1	—
Diazinon (4)	—	—
Parathion (5)	—	—

— no reaction observed. Values are the mean and standard deviation (\pm S.D.) of three independent experiments.

The reactivity order found is interesting. When piperidine is the nucleophile, malathion reacts faster than fenitrothion; both pesticides have thiophosphoryl moieties, but malathion has a better leaving group than fenitrothion. The lack of nucleophilic attack of piperidine on the electrophilic centers of paraoxon, parathion and diazinon in propylene carbonate as the solvent could be explained by the lower reactivity at the aliphatic carbon by the stereoelectronic effects of the ethoxy substituent.

Taking in account that parathion and diazinon did not react at room temperature, the degradation of these pesticides was studied under MW irradiation increasing the temperature in the range of 50–120 °C for 30 minutes using Bmim[Ala] as the nucleophile.

As shown in Fig. 1, at higher temperature, the extent of conversion (%) of both pesticides increases. Parathion's reaction with Bmim[Ala] is faster than with diazinon, and 100% of parathion's degradation occurs around 110 °C while at the same temperature diazinon degradation reaches 90%. Although the structures of both pesticides have certain similarities, such as an electrophilic P=S centre linked to -OEt groups, the aromatic substituents, *O*-phenyl in parathion and *O*-pyrimidyl in diazinon, can lead them to react through different reaction pathways (see below).

Subsequently, the MW-assisted nucleophilic degradation of paraoxon, parathion and diazinon with piperidine increasing the temperature in the range of 50–120 °C for 30 minutes was analyzed.

As we can see in Fig. 2, complete degradation of paraoxon requires higher temperature (120 °C) compared to parathion (90 °C) and diazinon (100 °C). This result is contrary to what was expected, because it has been described that P=O compounds are more reactive with nucleophiles than their thio-analogues.³⁷ This is an interesting finding and may provide a useful tool for the nucleophilic degradation of thiophosphate pesticides.

To evaluate the MW effect on the degradation of the pesticides studied, we carried out the same nucleophilic reactions of paraoxon, parathion and diazinon at 50 °C for 30 minutes



Fig. 1 Influence of MW heating on the degradation of parathion (5) (gray) and diazinon (4) (white) with Bmim[Ala] as the nucleophile and PC as the solvent at 50–120 °C for 30 minutes. Each experiment was performed in triplicate.

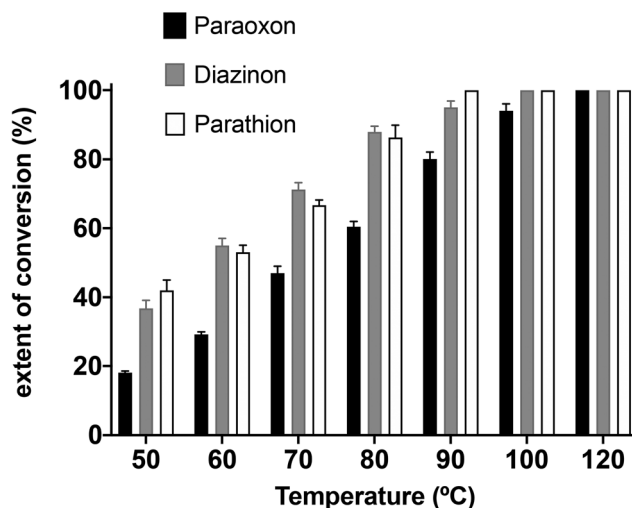


Fig. 2 Influence of MW heating on the degradation of parathion (5) (white), diazinon (4) (grey) and paraoxon (3) (black) with piperidine as the nucleophile at 50–120 °C for 30 minutes. Each experiment was performed in triplicate.

under conventional heating (oil bath) with controlled stirring and silent conditions (no heating, no stirring). The extent of conversion (%) in all cases was calculated by integration of the ³¹P NMR signals of each pesticide and is shown in Fig. 3.

As we can see in Fig. 3, there is a great influence of MW irradiation on the rate of the degradation of organophosphate pesticides compared with conventional heating (oil bath). In the case of silent conditions (no heating, no stirring), no degradation was observed after 30 minutes of reaction, which

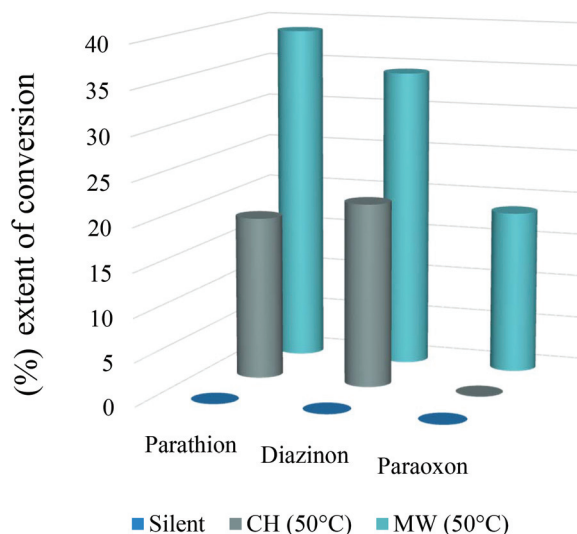


Fig. 3 Comparison of the extent of conversion (%) of pesticides after 30 minutes of reaction with piperidine as the nucleophile under MW heating (50 °C, light green blue columns), conventional heating (50 °C, dark green columns) and silent conditions (no heating, blue columns). Each experiment was performed in triplicate.

clearly demonstrates the effectiveness of the developed microwave process.

Keeping in mind that the reactivity of organophosphorus pesticides depends on their structure, the nature of the leaving group and nucleophile, and the solvent,²⁷ product analyses of the reactions were performed in order to understand the mechanism of the degradations. Then, the nucleophilic reactions were followed by the ³¹P NMR technique, and the spectra were recorded at different reaction times as shown in Fig. S1–S5 and S11–S13 in the ESI†

The highest degradation rate was found for the reaction of malathion with both nucleophiles at room temperature. This result is probably related to the very good electrophilic carbon center in malathion which makes it more reactive than the aromatic pesticides studied in this work. This finding is in agreement with a previous work of Lartiges *et al.* who studied the degradation kinetics of 19 organophosphorus and organo-nitrogen pesticides in different water types.³⁶ The ³¹P NMR spectrum for the degradation with piperidine showed two new phosphorus signals (Fig. S2 in the ESI†), one at 118 ppm and another at 67 ppm. The signal at 118 ppm was assigned to *O,O*-dimethyl dithiophosphate (**6**) which is produced by an S_N2 (C–S) pathway (aliphatic carbon bonded to a sulphur atom), identified by ESI-MS (–Q, *m/z* 156.97, Fig. S6 in the ESI†). In addition, the signal at 67 ppm in Fig. S2† was assigned to compound **8** (Scheme 2), which is formed when piperidine attacks the phosphorus center of malathion S_N2(P). The product **8** was identified by comparison with the phosphoric product

obtained by the reaction of piperidine and *O,O*-dimethyl chlorothiophosphate in PC, as shown in Scheme S1 and Fig. S7 in the ESI.† When the reaction of malathion with Bmim[Ala] at room temperature was followed by ³¹P NMR, only one new phosphorus signal (118.5 ppm) was observed. This was assigned to *O,O*-dimethyl dithiophosphate (**6**), produced by an S_N2(C–S) pathway.

Regarding the degradation of fenitrothion (**2**) at room temperature, the ³¹P NMR spectra showed only one phosphorus signal at 52.3 ppm when Bmim[Ala] was the nucleophile. This signal was attributed to demethylfenitrothion (**10**) formed when the nucleophile attacks the aliphatic carbon by an S_N2 (C) pathway (Fig. S1 in the ESI†); see Scheme 3. According to Fig. S4,† the nucleophilic degradation of fenitrothion (**2**) by piperidine in PC is similar and showed a phosphorus signal at 52 ppm, which was also assigned to demethylfenitrothion (**10**) as shown in Scheme 3.

On the other hand, the degradation of paraoxon (**3**) with Bmim[Ala] at room temperature shows the decrease of the signal of **3** (–7.4 ppm) and the increase of a phosphorus signal at –0.30 ppm (Fig. S3 in the ESI†), which was assigned to *O,O*-diethylphosphate (**12**); see Scheme 3. Compound **12** is formed when Bmim[Ala] attacks the aromatic carbon of paraoxon (S_NAr), involving an aryl–O breakage according to the literature.²⁷ Probably different interactions could explain these results: (i) stabilization of the transition state of the S_NAr reaction through a π stacking interaction between the phenyl moiety of paraoxon and the aromatic cation (Bmim) of the



Scheme 2 Reaction pathways for the nucleophilic degradation of malathion (**1**) with both nucleophiles (piperidine and Bmim[Ala]) in propylene carbonate at room temperature.



Scheme 3 Reaction pathways for the nucleophilic degradation of fenitrothion (**2**) by both nucleophiles (piperidine and Bmim[Ala]) and paraoxon (**3**) by Bmim[Ala] as the nucleophile in PC at room temperature.



Scheme 4 Reaction pathways for the nucleophilic degradation of diazinon (4) and parathion (5) by both nucleophiles (piperidine and Bmim[Ala]) in propylene carbonate under MW heating.

nucleophile and (ii) interaction of the imidazolium cation of the IL with the starting material paraoxon which was described by Hawker *et al.*³⁸ as an entropic effect.

Then, the product analysis of the nucleophilic degradation of the pesticides under MW heating in the range of 50–120 °C for 30 minutes was performed. In the reaction of diazinon with Bmim[Ala], a new signal appears at 49.5 ppm in the ³¹P NMR spectra (Fig. S8 in the ESI†), assigned to diethyl thiophosphate (14). This is produced when the nucleophile attacks the aromatic carbon of diazinon (S_NAr) and it was confirmed by ESI-MS (-Q, *m/z* 168.9) (Fig. S9 in the ESI†). The degradation of diazinon by piperidine after MW irradiation shows only one phosphoryl product (Fig. S10 in the ESI†), which is the same product 14 described above when Bmim[Ala] was the nucleophile (see Scheme 4).

In the case of parathion's degradation with Bmim[Ala], ³¹P NMR spectra at different times showed the decrease of the substrate signal (62 ppm) and the increase of a new signal at 50.5 ppm which was assigned to the *O*-ethyl 4-nitrothiophenylphosphate diester (16) (Fig. S11 in the ESI†). This product is formed by the nucleophilic attack of the nucleophile on the aliphatic carbon of parathion, through an S_N2(C) pathway as shown in Scheme 4. When the nucleophile is piperidine, it was seen that degradation occurred by the same pathway as described above (Fig. S12 in the ESI†), *i.e.*, *via* the attack on the aliphatic carbon of the ethyl group. The non-phosphoryl product of this reaction was identified by GC-MS and was found to be *N*-ethylpiperidine (17b) (*r.t.* = 17.15 min, *m/z* 113) (Fig. S13 in the ESI†).

A comparison of parathion and diazinon degradation under MW heating with both nucleophiles, piperidine and Bmim

[Ala], shows that in the whole range of studied temperatures, the variation in the extent of conversion (%) is very small. Therefore, we can choose Bmim[Ala] as the best nucleophile to degrade these pesticides under these experimental conditions.

Finally, degradation of paraoxon under MW heating using piperidine as the nucleophile shows only one phosphorus signal at -6 ppm (Fig. S14 in the ESI†), which was assigned to the *O*-ethyl 4-nitrophenylphosphate diester (18), formed when piperidine attacks the aliphatic carbon of paraoxon.²⁷ Nevertheless, when the reaction of paraoxon with piperidine was carried out using ionic liquids and bio-based solvents, the degradation process was not selective and three reaction pathways were observed: to the aliphatic carbon of *O*-ethyl, to the aromatic carbon C1 and to the phosphorus centre.²⁸ Interestingly, in this work, the microwave-assisted degradation of paraoxon and parathion in PC is selective and proceeds only through nucleophilic attacks by piperidine on the aliphatic carbon (S_N2(C)) of the pesticides resulting in products 16 and 18 (Schemes 4 and 5).

As can be seen, the aromatic OPP studied in this work (2–5) react solely *via* S_N2 at the aliphatic carbon when piperidine is the nucleophile. The preference for this pathway depends on the nucleophile^{39,40} but also depends on the solvent. It has been described that hard nucleophiles (oximates and ⁻OH) react with the hard P center and soft nucleophiles (butylamine) react with the soft aliphatic carbon center preferentially.^{39,41} Furthermore, the preference by the S_N2(C) in OPP's degradation has been reported before and it was explained as a consequence of the lower basicity of the medium.^{42,43}



Scheme 5 Reaction pathway for the degradation of paraoxon by piperidine as the nucleophile in propylene carbonate under MW heating.

Interestingly, the use of propylene carbonate is important to control the selectivity of this reaction. Previously, we have reported the degradation of paraoxon with piperidine in some conventional organic solvents and the $S_N2(C)$ pathway is the main degradation route (above 75%) in solvents less polar than DMSO, such as 1,4-dioxane and acetonitrile.²⁷ The k_{obsd} value found for the degradation of paraoxon in PC with Bmim [Ala] as the nucleophile (this study) is similar ($9.6 \times 10^{-6} \text{ s}^{-1}$) to those reported for the degradation of paraoxon in conventional ionic liquids such as Bmim[NTF₂], B₂mim[NTF₂] and B₂mim[BF₄] by the same reaction pathway.²⁷ On the other hand, in a previous study, the degradation of fenitrothion with piperidine was carried out in some bio-based solvents and conventional ionic liquids and the only reaction pathway found was the $S_N2(C)$ pathway, propylene carbonate being the best solvent to degrade this pesticide.²⁸ Finally, for malathion degradation, this study showed that the $t_{1/2}$ value was around 10 minutes with both nucleophiles, which is a very promising result since in previous works the degradation of malathion was studied in aqueous solutions using strong chemical oxidants and UV irradiation. The $t_{1/2}$ values were higher than 12 hours while the best method to degrade this pesticide was under O₃ radiation with a $t_{1/2}$ value of 2.4 hours.⁴⁴

These results are interesting from the toxicity point of view since it is known that phosphate monoesters and diesters are much less lipophilic, and therefore less toxic, than their triester precursor, paraoxon or parathion. It is important to mention that even at long reaction times under MW heating, the same products of the reaction were found in all cases.

4. Conclusions

An efficient process for the chemical degradation of organophosphorus pesticides through microwave-assisted degradation in the green solvent "propylene carbonate" was developed.

This study helps to understand the mechanism of the reaction of OPP which can be used to create efficient and sustainable degradation methods.

Fenitrothion and malathion do not need MW irradiation to achieve complete degradation with either nucleophile used in this study. Fenitrothion degradation was faster with piperidine as the nucleophile, while malathion degradation was carried out at a similar rate with both nucleophiles ($t_{1/2}$ values are 9.57 min and 10.9 min with piperidine and Bmim[Ala] respectively). On the other hand, the degradation of paraoxon with Bmim[Ala] was accomplished at room temperature in a few hours, which is an important result because the degradation with piperidine was not successful.

MW heating is a promising technique that accelerates the degradation of the OPP paraoxon, diazinon, and parathion with piperidine which is effective in only 30 minutes at above 90 °C.

The temperature does not affect the reaction pathways of the degradation reactions studied. For the reaction of para-

oxon, only one phosphorylic compound was found in contrast to the three phosphorylic compounds observed in the nucleophilic degradation of paraoxon in other solvents.²⁷

This study highlights the efficiency of the microwave-assisted degradation of OPP in propylene carbonate that proceeded at a significantly higher rate compared to conventional heating and silent conditions.

Conflicts of interest

There are no conflicts to declare.

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