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Meta-analysis of neonicotinoid insecticides in global surface waters

Jianqiao Wang,^{1†} Ru Yin,^{1†} Yilin Liu,¹ Beijia Wang,¹ Nana Wang,¹ Pengfei Xiao,² Tangfu Xiao,^{1,3} Hirofumi Hirai^{4,5*}

¹ Key Laboratory for Water Quality and Conservation of the Pearl River Delta, Ministry of Education, School of Environmental Science and Engineering, Guangzhou University, Guangzhou 510006, China

² College of Forestry, Northeast Forestry University, Harbin 150040, China

³ State Key Laboratory of Geohazard Prevention and Geoenvironment Protection, Chengdu University of Technology, Chengdu 610059, China

⁴ Faculty of Agriculture, Shizuoka University, 836 Ohya, Suruga-ku, Shizuoka 422-8529, Japan

⁵ Research Institute of Green Science and Technology, Shizuoka University, 836 Ohya, Suruga-ku, Shizuoka 422-8529, Japan

† Jianqiao Wang and Ru Yin made equal contribution to this work.

* Corresponding author: Hirofumi Hirai

Mailing address: Faculty of Agriculture, Shizuoka University, 836 Ohya, Suruga-ku, Shizuoka 422-8529, Japan

Tel. & Fax: +81 54 238 4853

E-mail address: hirai.hirofumi@shizuoka.ac.jp

Abstract

1 Neonicotinoids (NEOs) are a class of insecticides that have high insecticidal activity and
2 are extensively used worldwide. However, increasing evidence suggests their long-term
3 residues in the environment and toxic effects on nontarget organisms. NEO residues are
4 frequently detected in water and consequently have created increasing levels of pollution and
5 pose significant risks to humans. Many studies have focused on NEO concentrations in water;
6 however, few studies have focused on global systematic reviews or meta-analyses of NEO
7 concentrations in water. The purpose of this review is to conduct a meta-analysis on the
8 concentration of NEOs in global waters based on published detections from several countries
9 to extend knowledge on the application of NEOs. In the present study, 43 published papers
10 from 10 countries were indexed for a meta-analysis of the global NEO distribution in water.
11 Most of these studies focus on the intensive agricultural area, such as eastern Asia and North
12 America. The order of mean concentrations is identified as imidacloprid ($119.542 \pm 15.656 \text{ ng}$
13 L^{-1}) > nitenpyram ($88.076 \pm 27.144 \text{ ng L}^{-1}$) > thiamethoxam ($59.752 \pm 9.068 \text{ ng L}^{-1}$) >
14 dinotefuran ($31.086 \pm 9.275 \text{ ng L}^{-1}$) > imidaclothiz ($24.542 \pm 2.906 \text{ ng L}^{-1}$) > acetamiprid
15 ($23.360 \pm 4.015 \text{ ng L}^{-1}$) > thiacloprid ($11.493 \pm 5.095 \text{ ng L}^{-1}$). Moreover, the relationships
16 between NEO concentrations and some environmental factors is analyzed. NEO
17 concentrations increase with temperature, oxidation-reduction potential and the percentage of
18 cultivated crops but decrease with stream discharge, pH, dissolved oxygen, and precipitation.
19 NEO concentrations show no significant relations to turbidity and conductivity.

Keywords: Neonicotinoids; insecticide; organic pollutant; insecticide contamination; water;
meta-analysis

1. Introduction

20 Neonicotinoids (NEOs) are a class of insecticides that act selectively on nicotinic
21 acetylcholine receptors (nAChRs) to block the action of acetylcholine in the central nervous
22 systems of insects (Matsuda et al. 2001; Tomizawa and Casida 2003). Compared to traditional
23 pesticides, they show stronger selectivity for insects on nAChRs than vertebrates and are thus
24 considered to have reduced toxicity and to exhibit lower resistance in mammals (Jeschke et al.
25 2013). Since NEOs were first produced in the 1990s beginning with imidacloprid (IMI), other
26 NEOs, including acetamiprid (ACE), clothianidin (CLO), thiamethoxam (TXM), thiacloprid
27 (THI), nitenpyram (NIT), and dinotefuran (DIN), have been successively developed for the
28 market (Godfray et al. 2015). In addition, imidaclothiz (IMZ) is a new NEO with more
29 systemic activity developed by Nantong Jiangshan Agrochemical and Chemical Co. Ltd.,
30 China and it was registered in 2006 by the Chinese Ministry of Agriculture (Shao et al. 2013).
31 NEOs have become best-selling insecticides with annual sales of 1.9 billion dollars,
32 accounting for 25% of the global insecticide market since 2010 (Jeschke et al. 2011). In 2012,
33 TXM, CLO, and IMI accounted for almost 85% of total NEO sales and were mainly used for
34 crop protection (Bass et al. 2015). In particular, IMI has gradually become one of the most
35 widely applied insecticides and is used for over 140 agricultural crops in approximately 120
36 countries (Drobne et al. 2008). Approximately 20,000 tons of active substance IMI is
37 produced annually, and China contributes approximately 70% of IMI production (Drobne et
38 al. 2008; Simon-Delso et al. 2015; Wang et al. 2018). Because of the highly efficient insect
39 pest control and favorable safety profiles of NEOs, they have been used in agriculture, animal
40 husbandry, and residential environments worldwide (Simon-Delso et al. 2015; Morrissey et
41 al. 2015).

42 Along with their global use, NEOs have had negative effects on wildlife. Many
43 organisms, including nontarget species and terrestrial pollinators such as bumble bee (*Bombus*
44 *terrestris*), honey bee (*Apis mellifera*), and butterfly (*Polyommatus icarus*), are extremely
45 sensitive to NEOs (Whitehorn et al. 2012; Rundlöf et al. 2015; Basley and Goulson 2018).
46 Honey bees, as pollinators, play essential roles in ecological systems and crop productivity, so

47 their health, productivity and behavior are of greater environmental concern (Henry et al.
48 2012). An increasing number of studies have revealed that NEOs tend to easily enter
49 ecosystems through runoff and drainage systems in agricultural areas and pose increasing
50 ecological threats to organisms (Anderson et al. 2018; Schaafsma et al. 2019). NEOs have the
51 potential to cause a sudden decline in the adult honeybee population, also known as colony
52 collapse disorder (Henry et al. 2012). Many studies have reported on the acute toxicity of
53 NEOs to aquatic invertebrates, birds, and mammals from *in vitro* and *in vivo* laboratory
54 toxicity experiments (Morrissey et al. 2015; Han et al. 2018; Addy-Orduna et al. 2019). The
55 potential toxic effects of NEOs mainly include reproductive toxicology, neurotoxicity,
56 hepatotoxicity, immunotoxicity, and genetic toxicity (Han et al. 2018).

57 Variable levels of NEOs and their metabolites occur in surface environmental media
58 such as soils (Jones et al. 2014; Bonmatin et al. 2019), drinking water (Sultana, et al. 2018),
59 crops (Kamel et al. 2010; Chahil et al. 2015; Karthikeyan et al. 2019), pollen (Tosi et al.
60 2018), and even bovine milk (Adelantado et al. 2018). It is important to develop better
61 knowledge of the distribution of NEO levels in the environment and the associated
62 environmental effects, which will help guide conservation efforts to NEOs application and
63 environment protection. Meta-analysis is a quantitative method to summarize the independent
64 research results. Hence, the objective of this review is to summarize the global concentration
65 distribution of NEOs (ACE, CLO, DIN, IMI, IMZ, NIT, THI, and TXM) in water and reveal
66 the relationship between NEO concentrations and hydrologic parameters such as stream
67 discharge, turbidity, pH, temperature, dissolved oxygen (DO), oxidation-reduction potential
68 (ORP), precipitation, and cultivated crops via meta-analysis.

2. Materials and methods

69 2.1 Data assembly

70 To study NEO levels in water, target publications included in the PubMed database were
71 screened on February 2, 2021. A total of 57 papers were obtained using the following search
72 terms: (((neonicotinoid[Title]) OR (neonicotinoids[Title]) OR (neonicotinoid
73 insecticide[Title]) OR (neonicotinoid insecticides[Title])) AND ((water[Title]) OR

74 (lake[Title]) OR (river[Title]) OR (stream[Title]) OR (wetland[Title])). Among the papers
75 obtained, 27 were retained in the present study based on the following criteria: (1) papers
76 written in English were retained; (2) duplicate papers were removed; (3) irrelevant papers
77 were carefully removed after reading the abstracts; (4) papers excluding NEO concentration
78 data were removed after reading the full text in detail; and (5) papers were identified as
79 original research rather than review articles. An additional 16 papers were obtained from the
80 references of the retained papers, so a total of 43 papers were used in this study. These selected
81 papers were published from 2012 to 2021 with the impact factor range from 1.755 to 11.236.
82 Although they might be not comprehensive, the papers that we screened were published in
83 specialized journals with considerable impact. The following information was extracted:
84 sampling time, country, sampling location, physical and chemical properties of the studied
85 water (stream discharge, turbidity, pH, temperature, DO, ORP, and conductivity),
86 precipitation, percentage of cultivated crops, types of NEOs, concentrations of NEOs
87 (maximum, median, minimum, and mean), and standard deviation of NEO concentrations.
88 These studies referring to 10 countries (the United States, Australia, Belize, Canada, China,
89 Japan, the Philippines, Romania, South Africa, and Vietnam) were selected. NEOs were
90 detected in tap water, seawater, lakes, rivers, reservoirs, estuaries, creeks, wetlands, or open
91 ditches and runoff in agricultural regions whether it's spring, summer, fall or winter (Table
92 S1). Plot Digitizer software was used to extract values from graphs.

93 **2.2 Data analysis**

94 The sampling locations were displaced on a world map based on longitude and latitude
95 parameters by RStudio (Fig. 1). With no information on longitude and latitude, the sampling
96 site name was used to extract longitude and latitude information from Google Maps. The
97 mean concentration of each NEO was used, and the concentrations of NEOs were unified to
98 ng L⁻¹ for further analysis. Data analyses and the meta-analysis figures were developed using
99 the JMP statistical program (version 16.0). JMP is a statistical visualization tool, it can
100 integrate the graphics into the report. The “Distribution of Y” platform was used for testing
101 the mean concentrations of different NEOs. The number of observations and concentration

102 range for different NEOs (ACE, CLO, DIN, IMI, IMZ, NIT, THI, and TXM) were
103 summarized. The “Fit Y by X” platform was used for testing the significant differences
104 between the mean concentrations of NEOs and environmental factors (e.g., stream discharge,
105 turbidity, pH, temperature, DO, ORP, conductivity, precipitation, and the percentage of
106 cultivated crops).

3. Results and discussion

107 3.1 Database availability

108 The main regions exhibiting NEO use in agriculture are 29.4% of total global use in
109 Latin America, followed by 23% in Asia, North 22% in America, and 11% in Europe (Bass et
110 al. 2015; Simon-Delso et al. 2015). Most of our selected studies focus on eastern Asia and
111 North America, which include countries heavily focused on agricultural production (Fig. 1).
112 However, no study about Latin America was obtained in the present study. The mean
113 concentrations of eight widely used NEOs (ACE, CLO, DIN, IMI, IMZ, NIT, THI, and TXM)
114 were collected, and the information on each form of NEO detected is shown in Fig. 2. IMI is
115 the most frequently reported (39/43, 91%), followed by CLO (36/43, 84%), TXM (32/43,
116 74%), ACE (31/43, 72%), THI (27/43, 63%), DIN (16/43, 37%), NIT (11/43, 26%), and IMZ
117 (4/43, 9%). IMI, the first NEO developed, is the most frequently reported, possibly due to its
118 broad application and usage (Kollmeyer et al. 1999). IMZ was the latest to enter the market,
119 thus there only a few studies include IMZ detection. Continuous detection of IMZ in the
120 environment is necessary, because it has great potential in China's market.

121 3.2 NEO concentrations in water

122 Table 1 shows the concentrations and numbers of observations for different NEOs. CLO
123 was the most frequently detected in 1056 out of 1645 water samples, followed by IMI (879),
124 TXM (863), ACE (428), THI (295), DIN (122), IMZ (37), and NIT (29). CLO has the highest
125 mean concentrations at $222.320 \pm 46.692 \text{ ng L}^{-1}$. The mean concentrations of other NEOs are
126 ordered as follows: IMI ($119.542 \pm 15.656 \text{ ng L}^{-1}$) > NIT ($88.076 \pm 27.144 \text{ ng L}^{-1}$) > TXM

127 $(59.752 \pm 9.068 \text{ ng L}^{-1}) > \text{DIN} (31.086 \pm 9.275 \text{ ng L}^{-1}) > \text{IMZ} (24.542 \pm 2.906 \text{ ng L}^{-1}) > \text{ACE}$
128 $(23.360 \pm 4.015 \text{ ng L}^{-1}) > \text{THI} (11.493 \pm 5.095 \text{ ng L}^{-1})$. Moreover, concentrations were found
129 to range from 0.001 to 45100 ng L^{-1} for CLO, from 0.004 to 9140 ng L^{-1} for IMI, from 0.002
130 to 4315 ng L^{-1} for TXM, from 0.002 to 3820 ng L^{-1} for ACE, from 0.003 to 1370 ng L^{-1} for
131 THI, from 0.11 to 1022.2 ng L^{-1} for DIN, from 2 to 672.9 ng L^{-1} for NIT, and from 0.002 to
132 81.92 ng L^{-1} for IMZ (Table 1).

133 Fig. 3 displays the distributions of the mean concentrations of each NEO type. The
134 concentrations of CLO and IMI were found to be concentrated at 0~1500 ng L^{-1} and 0~500 ng
135 L^{-1} , respectively. The concentrations of ACE, DIN, IMZ, NIT, THI, and TXM were mainly
136 measured at below 250 ng L^{-1} . NEOs can be used in pest control to protect crops and are
137 mainly applied for seed treatment, chemigation, and soil treatment (Simon-Delso et al. 2015).
138 NEOs may enter through various media into aquatic systems from agricultural fields through
139 processes such as spray drift, atmospheric deposition, soil erosion, and runoff. Some
140 governments and organizations have established water quality guidelines for protecting
141 aquatic ecosystems. For example, the United States Environmental Protection Agency
142 (USEPA) has estimated that chronic benchmarks of 970, 2100, 10, 740, 95300, and 50 ng L^{-1}
143 for THI, ACE, IMI, TXM, DIN, and CLO, respectively (USEPA, 2016). In this review, some
144 potentially threatening concentrations of certain NEOs are especially found in agricultural
145 regions. THI monitored at the outlet of the Yarramundi Lagoon in a turf farm was found the
146 highest concentration of 1370 ng L^{-1} (Sánchez-Bayo et al. 2014). The highest IMI
147 concentration found in Solomon Creek in the Californian agricultural region was recorded as
148 9140 ng L^{-1} (Anderson et al. 2018). Although the province of Ontario of Canada bans the
149 cosmetic use of some pesticides on lawns and gardens, NEOs are used for seed treatment on
150 row crops such as corn, soybeans, cereal grains, and canola, which has led to widespread use
151 in Ontario (Ontario 2016). CLO, TXM, and ACE levels in drain water around maize fields in
152 Canada have reached 45100 and 7200, 4315, and 1527.6 ng L^{-1} , respectively (Schaafsma et al.
153 2019).

154 In recent years, the European Union has banned some NEOs because of their
155 improvement in the decline of bees and other pollinators (Naumann et al. 2022). However,
156 NEOs are still widely used in developing countries with poorly controlled. China has the
157 highest production of NEOs, which are frequently detected in rivers flowing through urban
158 environments. In addition to those found in agricultural regions, the highest concentrations of
159 DIN, NIT, and IMZ have been detected in the Yangtze River in China, reaching levels of
160 1022.3, 672.9, and 81.92 ng L⁻¹, respectively (Chen et al. 2019). The Yangtze River is the
161 longest river in China, playing a considerable role in agricultural and industrial activities
162 (Mahai et al. 2019). NEOs in the Yangtze River have become a source of NEOs in seawater
163 (Chen et al. 2019). Although NEO concentrations decrease rapidly by dilution, NEOs are
164 detected near shorelines (Pan et al. 2020). IMZ is a novel NEO that has been gradually
165 applied to vegetables, fruits, and crops on a large scale in China because of its excellent
166 insecticidal activity (Tao et al. 2021). Due to IMZ's increasing use, more attention should be
167 dedicated to its adverse effects (e.g., DNA damage in earthworms; Zhang et al. 2017).
168 Moreover, different NEO concentrations have been detected in different crop planting periods.
169 Concentrations of IMI and TXM increase markedly in the rice planting month. DIN was
170 detected at a concentration of 220 ng L⁻¹ during rice earwig emergence (Yamamoto et al.
171 2012). A large proportion of pesticides enter environmental media via runoff, leaching, and
172 drifting. These pesticides are absorbed by nontarget plants or organisms and present a
173 potential threat to food safety (Li et al. 2018; Tao et al. 2021). Thus, scientists around the
174 world have gradually recognized NEO risks and increased efforts to monitor NEOs in the
175 environment (Morrissey et al. 2015).

176 **3.3 Effect of physicochemical properties on NEO concentration**

177 Fig. 4 and Table 2 present the relationship between NEO concentrations and nine
178 physical and chemical properties. Different properties show different responses to NEO
179 concentrations in water. NEO concentrations increase with temperature, ORP, and the
180 percentage of cultivated crops (Line regression, Temperature: adjusted R² = 0.0811, *p* <
181 0.0001; ORP: adjusted R² = 0.0931, *p* < 0.01; Cultivated crop: adjusted R² = 0.0307, *p* <

182 0.001) (Fig. 4d, f, i). When summer arrives, pest damage increases with increasing
183 temperature, and insecticide use is increased to decrease crop losses. Rainfall is a key factor
184 in increasing NEO residues in water. NEOs can enter water via surface and underground
185 runoff, creating higher insecticide concentrations in water. For instance, in the province of
186 Guangdong located in the subtropical zone of South China, the climate is warm and humid for
187 most of the year. Thus, large quantities of pesticides are used for pest control, and Guangdong
188 Province has the highest pesticide application dosage (Li et al. 2014). Only one paper presents
189 the value of ORP, and the representativeness of the relation needs to be further confirmed (Yi
190 et al. 2019). Concentrations of NEOs generally increase as the percentage of cultivated crops
191 increases. High NEO concentrations are detected in surface water around areas of agricultural
192 activity when the planting season arrives. According to a study conducted in the USA, streams
193 show higher NEO concentrations in the planting season than in other seasons (Hladik and
194 Kolpin 2016). Another study from Canada shows that one side of the Two Mile Creek
195 watershed includes over 50% orchards, and an IMI concentration of 816 ng L⁻¹ was detected
196 in this creek (Struger et al. 2017). A positive relationship between cultivated crops and NEO
197 concentrations have been observed in other studies (Hladik et al. 2014; Iancu et al. 2019).

198 NEO concentrations decrease with stream discharge, pH, DO, and precipitation (Line
199 regression, Stream discharge: adjusted R²=0.0433, $p > 0.05$; pH: adjusted R² = 0.0225, $p <$
200 0.01; DO: adjusted R² = 0.0794, $p < 0.01$; Precipitation: adjusted R² = 0.0223, $p < 0.0001$)
201 (Fig. 4a, c, e, g). The negative relation between NEO concentrations and stream discharge or
202 precipitation may be caused by the dilution of NEOs when strong precipitation occurs
203 (Struger et al. 2017). Higher DO value of water might affect the degradation of NEOs (Yi et
204 al. 2019). The pH value is an important factor that affects NEO solubility in water. NEOs
205 have longer term residuals under acidic, or neutral conditions than under less alkaline
206 conditions (Yi et al. 2019). It was reported that NEOs hardly degrade at pH 4.0 ~ 7.0, while
207 NEOs hydrolyze readily with a high pH value (pH = 10). (Todey et al. 2018). In this review,
208 pH values of water samples were ranged from 6.31 to 8.67, suggesting that NEOs might be
209 presented in waters for a long time.

210 The NEO concentrations show no significant correlations with turbidity, and
211 conductivity ($p > 0.05$) (turbidity: adjusted $R^2 = -0.00781$, $p = 0.879$; conductivity: adjusted
212 $R^2 = 0.00456$, $p = 0.184$) (Fig. 4b, h). NEOs are more likely to dissolve than combine with
213 particulate, or colloidal matter (Sánchez-Bayo and Hyne 2014). However, these relationships
214 need further confirmation.

4. Conclusions and avenues for future research

215 In the present work, we summarize a total of 43 publications on NEOs detected in tap
216 water, seawater, lakes, rivers, reservoirs, estuaries, creeks, wetlands, open ditches, and runoff
217 in agricultural regions worldwide. Most studies have focused on eastern Asia and North
218 America, which are major areas of agricultural production. The order of reporting frequency
219 is $IMI > CLO > TXM > ACE > THI > DIN > NIT > IMZ$. Underdeveloped areas such as
220 Africa should be considered due to an increasing use of NEOs in these areas. In addition, the
221 order of mean concentrations is $IMI > NIT > TXM > DIN > IMZ > ACE > THI$. The highest
222 IMI concentration (9140 ng L^{-1}) was detected in Solomon Creek in the Californian
223 agricultural region of the USA, while THI (1370 ng L^{-1}) was monitored at the outlet of the
224 Yarramundi Lagoon in Australia. The highest concentrations of CLO (45100 ng L^{-1} , 7200 ng
225 L^{-1}), TXM (4315 ng L^{-1}) and ACE (1527.6 ng L^{-1}) were found in drain water around maize
226 fields in Canada, and DIN (1022.3 ng L^{-1}), NIT (672.9 ng L^{-1}), and IMZ (81.92 ng L^{-1}) were
227 detected in the Yangtze River in China. Moreover, the relationships between mean
228 concentrations of NEOs and environmental factors (e.g., stream discharge, turbidity, pH,
229 temperature, DO, ORP, conductivity, precipitation, and the percentage of cultivated crops)
230 show that NEO concentrations increase with temperature, oxidation-reduction potential, and
231 the percentage of cultivated crops but decrease with stream discharge, pH, DO and
232 precipitation. NEO concentrations have no significant relationship to turbidity, and
233 conductivity. To prevent NEO pollution, NEO levels in the environment should be constantly
234 monitored and evaluated.

235

236 **Abbreviations**

237 **NEOs**, Neonicotinoids; **ACE**, acetamiprid; **CLO**, clothianidin; **DIN**, dinotefuran; **IMI**,
238 imidacloprid; **IMZ**, imidaclothiz; **NIT**, nitenpyram; **THI**, thiacloprid; **TXM**, thiamethoxam;
239 **DO**, dissolved oxygen, **ORP**, oxidation-reduction potential; **ND**, not detected; **NA**, not
240 analyzed; **MDL**, method detection limit.

241

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257 **Ethical Approval:** There are no ethical issues in this article.

258 **Consent to Participate:** All the authors agree to participate in this paper.

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260

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420 **Figures and Tables Captions**

421 Fig. 1 Geographic focuses of field studies investigating concentrations of NEOs in water
422 worldwide.

423 Fig. 2 Number of papers focused on each NEO concentration in water.

424 Fig. 3 Distribution of mean concentrations of each NEO (a: ACE; b: CLO; c: DIN; d: IMI; e:
425 IMZ.; f: NIT; g: THI; h: TXM). The top and bottom of the diamond (graph on the right)
426 are a 95% confidence interval for the mean. The bottom and top of the box show the
427 25th and 75th quantiles, and median is the horizontal line inside the box.

428 Fig. 4 NEO concentration responses to the effects of stream discharge (a), turbidity (b), pH
429 (c), temperature (d), DO (e), ORP (f), precipitation (g), conductivity (h), and the percentage of
430 cultivated crops (i). $p < 0.05$: statistically significant change; $p < 0.001$: highly
431 statistically significant.

432 Table 1 Summary of the dataset indicating the number of observations for different NEO
433 types (ACE, CLO, DIN, IMI, DIN, IMZ, NIT, THI, TXM), and statistics (Mean \pm standard
434 error (SE), lower 95% confidence interval (LCI), upper 95% confidence interval (UCI)) and
435 the ranges of concentrations of each NEO type.

436 Table 2 Description of the models that explain the relationships between mean concentrations
437 of NEOs and stream discharge, turbidity, pH, temperature, dissolved oxygen, ORP,
438 precipitation, conductivity, and the percentage of cultivated crops.

439 **Supplementary data**

440 Table S1 Raw data.

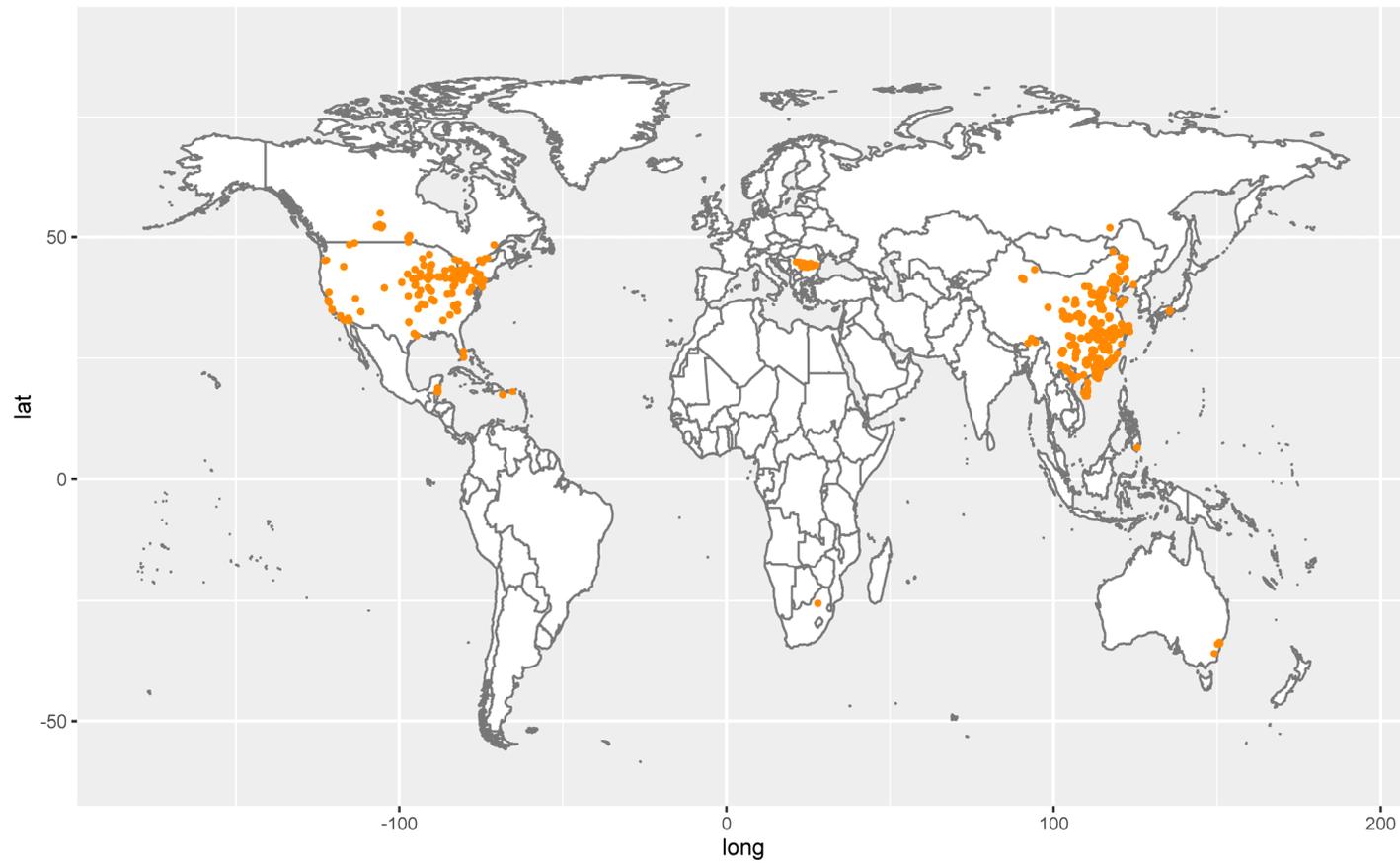


Fig. 1 Location of field studies that investigated the concentration of NEOs in water worldwide.

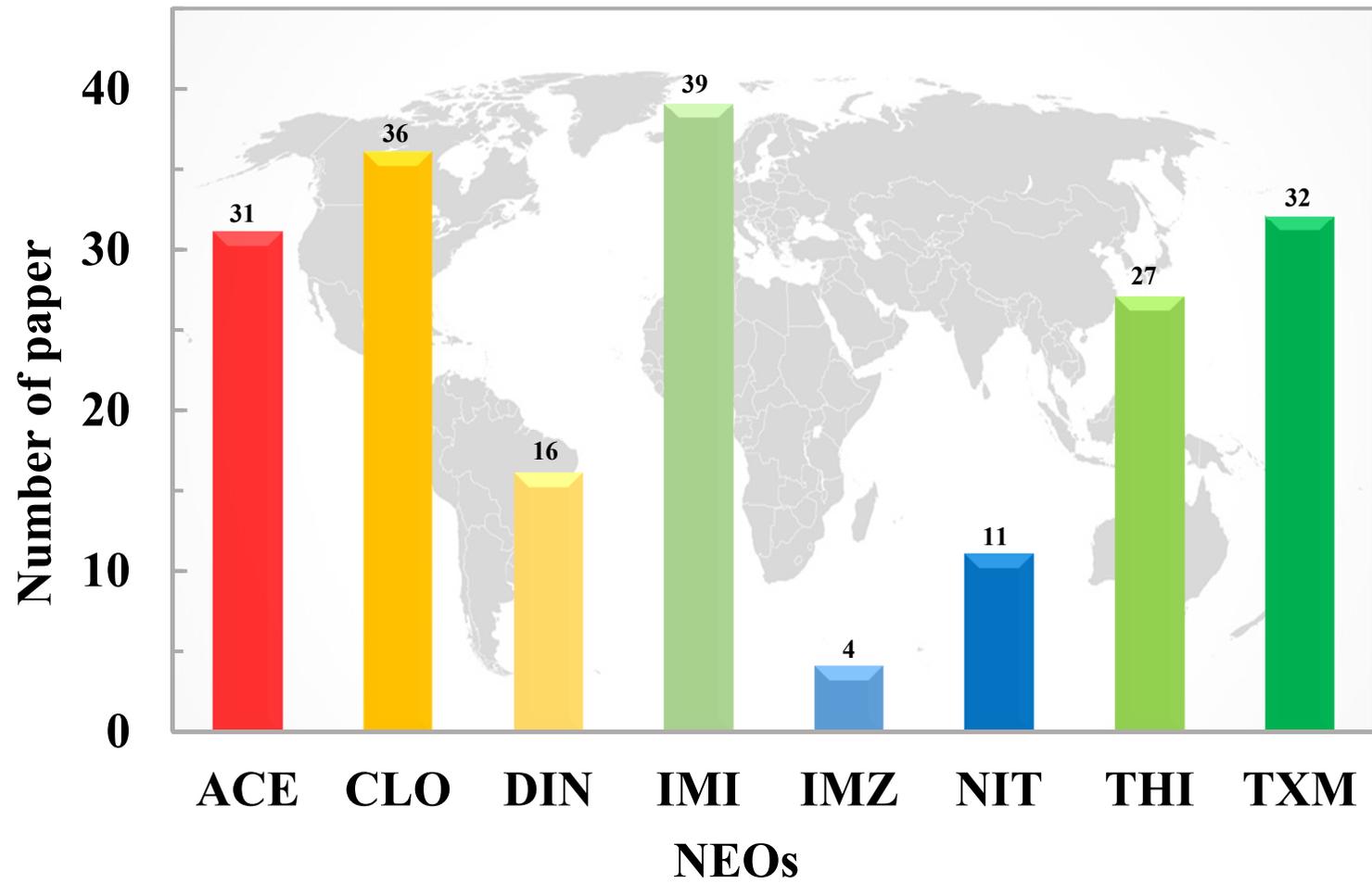


Fig. 2 The number of papers conducted on each NEOs concentration detection in waters.

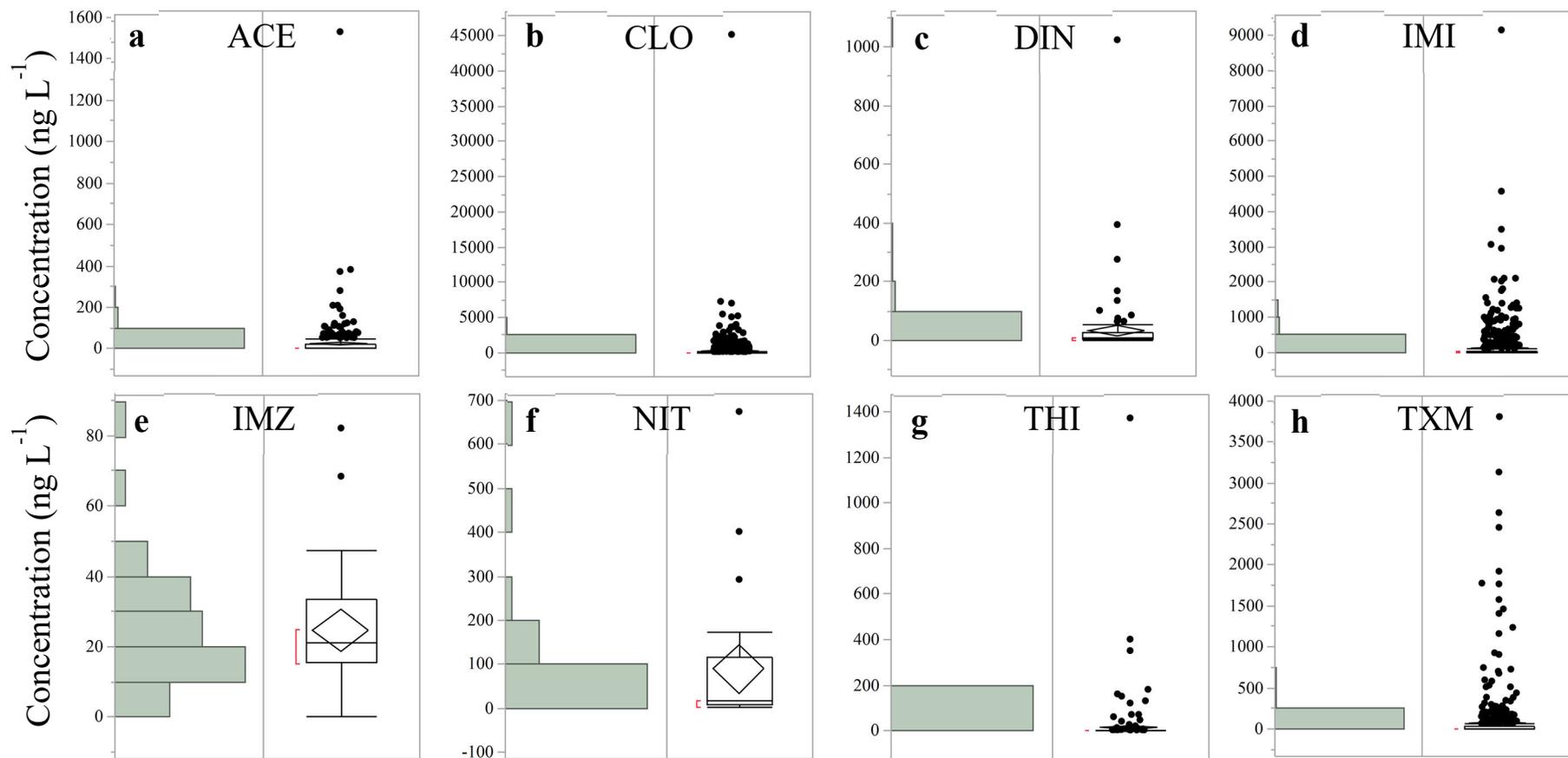


Fig. 3 Distribution of mean concentrations of each NEO (a: ACE; b: CLO; c: DIN; d: IMI; e: IMZ.; f: NIT; g: THI; h: TXM). The top and bottom of the diamond (graph on the right) are a 95% confidence interval for the mean. The bottom and top of the box show the 25th and 75th quantiles, and median is the horizontal line inside the box.

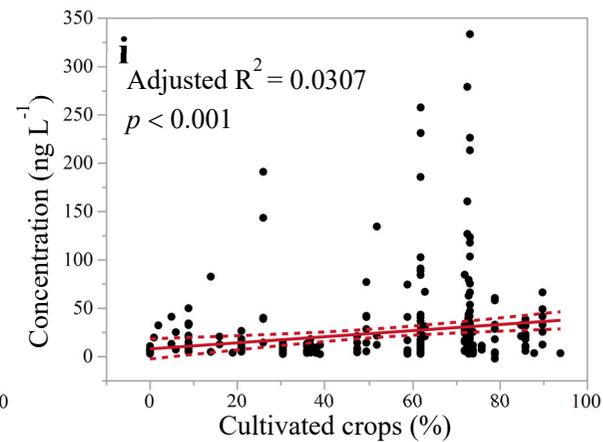
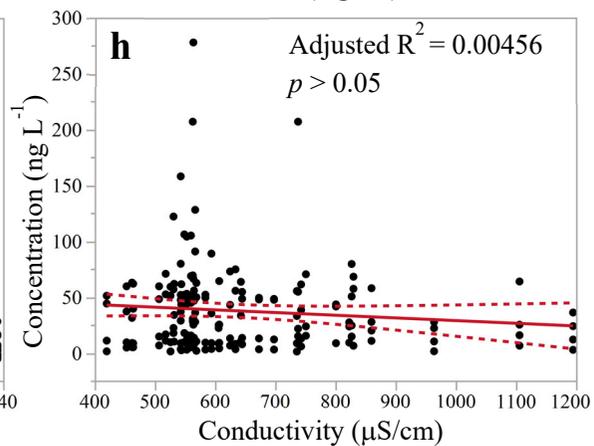
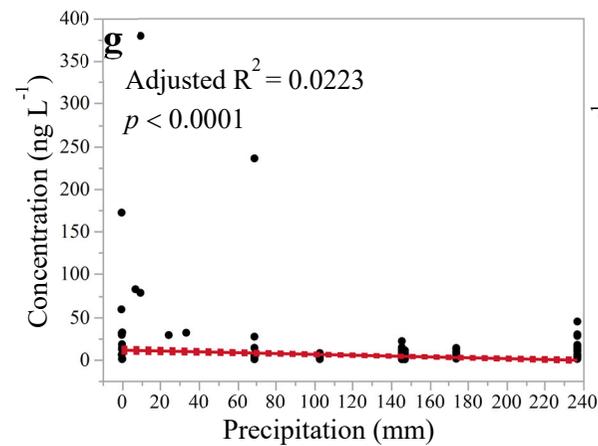
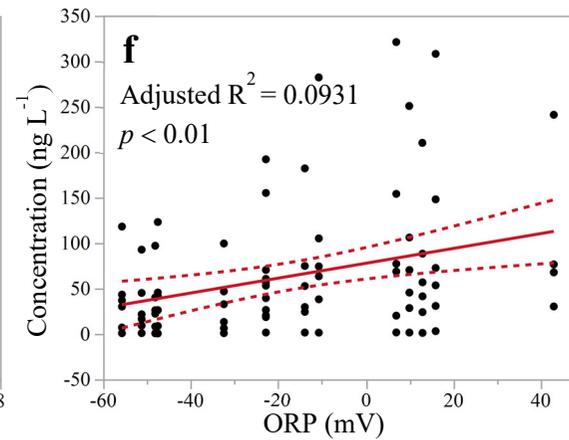
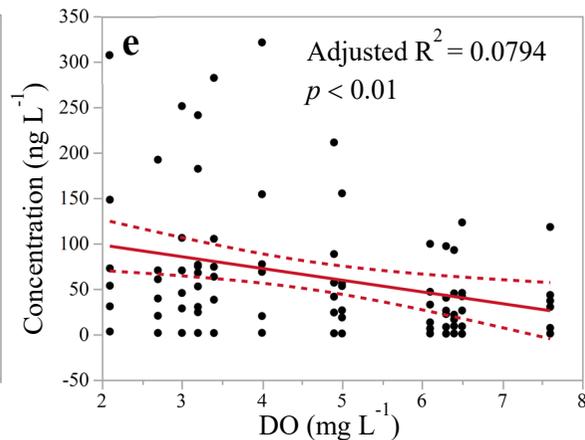
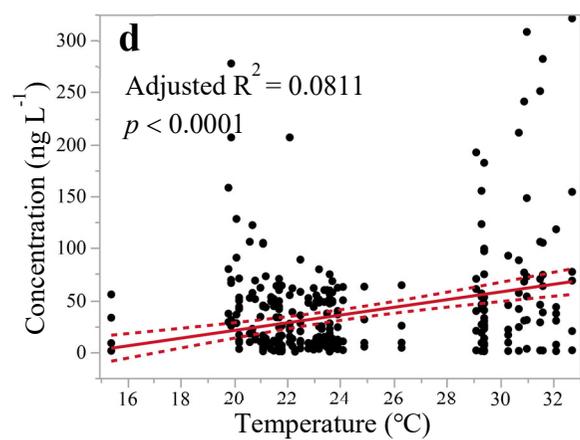
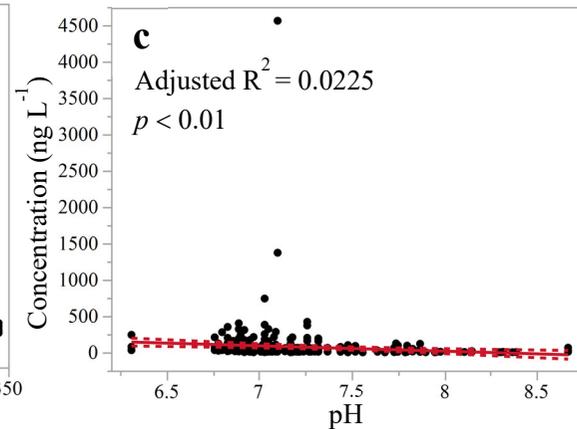
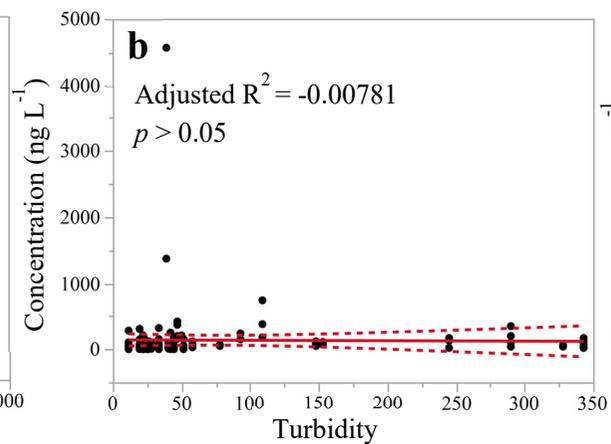
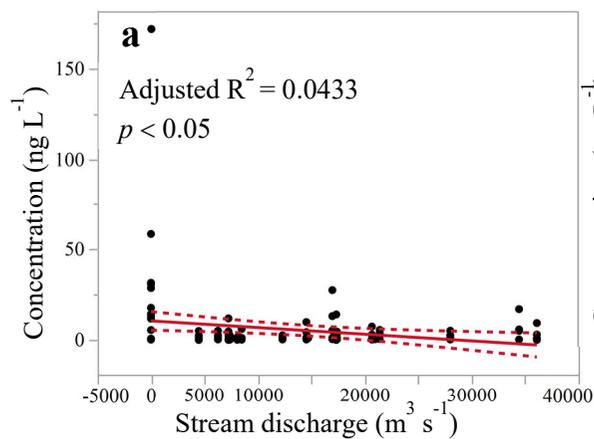


Fig. 4 NEO concentration responses to the effects of stream discharge (a), turbidity (b), pH (c), temperature (d), DO (e), ORP (f), precipitation (g), conductivity (h), and the percentage of cultivated crops (i). $p < 0.05$: statistically significant change; $p < 0.001$: highly statistically significant.

Table 1. Summary of the dataset indicating the number of observations for different NEO types (ACE, CLO, DIN, IMI, DIN, IMZ, NIT, THI, TXM), and statistics (Mean \pm standard error (SE), lower 95% confidence interval (LCI), upper 95% confidence interval (UCI)) and the ranges of concentrations of each NEO type.

| Type | n | Mean (ng L ⁻¹) | SE | Range (ng L ⁻¹) | LCI | UCI |
|------|------|----------------------------|--------|-----------------------------|---------|---------|
| ACE | 428 | 23.360 | 4.015 | [0.0025, 1527.6] | 15.469 | 31.252 |
| CLO | 1056 | 222.320 | 46.692 | [0.001, 45100] | 130.700 | 313.939 |
| DIN | 122 | 31.086 | 9.275 | [0.11, 1022.2] | 12.725 | 49.448 |
| IMI | 879 | 119.542 | 15.656 | [0.004, 9140] | 88.813 | 150.270 |
| IMZ | 37 | 24.542 | 2.906 | [0.002, 81.92] | 18.648 | 30.436 |
| NIT | 29 | 88.076 | 27.144 | [2, 672.9] | 32.475 | 143.678 |
| THI | 295 | 11.493 | 5.095 | [0.003, 1370] | 1.466 | 21.520 |
| TXM | 863 | 59.752 | 9.068 | [0.002, 3820] | 41.960 | 77.543 |

Table 2. Description of the models that explain the relationships between mean concentrations of NEOs and stream discharge, turbidity, pH, temperature, dissolved oxygen, ORP, precipitation, conductivity, and the percentage of cultivated crops.

| Model | R ² | Adjusted R ² | F value | <i>p</i> | n |
|---|----------------|-------------------------|----------------------------|----------|-----|
| Mean Concentration = 10.545 - 0.000368*Stream discharge | 0.0510 | 0.0433 | F _{1,125} =6.658 | 0.011 | 126 |
| Mean Concentration = 141.816 - 0.0639*Turbidity | 0.000187 | -0.00781 | F _{1,126} =0.0234 | 0.879 | 127 |
| Mean Concentration = 607.822 - 73.932*pH | 0.0248 | 0.0225 | F _{1,429} =10.872 | 0.0011 | 430 |
| Mean Concentration = -53.602 + 3.708*Temperature | 0.0839 | 0.0811 | F _{1,339} =30.954 | <0.0001 | 340 |
| Mean Concentration = 124.006 - 12.910*DO | 0.0906 | 0.0794 | F _{1,82} =8.0743 | 0.0057 | 83 |
| Mean Concentration = 77.593 + 0.817*ORP | 0.104 | 0.0931 | F _{1,82} =9.421 | 0.0029 | 83 |
| Mean Concentration = 10.796 - 0.0497*Precipitation | 0.0236 | 0.0223 | F _{1,734} =17.736 | <0.0001 | 735 |
| Mean Concentration = 52.817 - 0.024*Conductivity | 0.0104 | 0.00456 | F _{1,170} =1.778 | 0.184 | 171 |
| Mean Concentration = 7.237 + 0.314*Cultivated crops (%) | 0.0336 | 0.0307 | F _{1,331} =11.480 | 0.0008 | 332 |