

# Do estrogenic compounds in drinking water migrating from plastic pipe distribution system pose adverse effects to human? An analysis of scientific literature

Ze-hua Liu<sup>1,2,3</sup> · Hua Yin<sup>1</sup> · Zhi Dang<sup>1</sup>

Received: 8 June 2016 / Accepted: 1 November 2016 / Published online: 9 November 2016  
© Springer-Verlag Berlin Heidelberg 2016

**Abstract** With the widespread application of plastic pipes in drinking water distribution system, the effects of various leachable organic chemicals have been investigated and their occurrence in drinking water supplies is monitored. Most studies focus on the odor problems these substances may cause. This study investigates the potential endocrine disrupting effects of the migrating compound 2,4-di-tert-butylphenol (2,4-d-t-BP). The summarized results show that the migration of 2,4-d-t-BP from plastic pipes could result in chronic exposure and the migration levels varied greatly among different plastic pipe materials and manufacturing brands. Based on estrogen equivalent (EEQ), the migrating levels of the leachable compound 2,4-d-t-BP in most plastic pipes were relative low. However, the EEQ levels in drinking water migrating from four out of 15 pipes may pose significant adverse effects. With the increasingly strict requirements on regulation of drinking water quality, these results indicate that some drinking water transported with plastic pipes may not be safe for human consumption due to the occurrence of 2,4-d-t-BP. Moreover, 2,4-d-t-BP is not the only plastic pipe-migrating estrogenic compound, other compounds such as 2-

tert-butylphenol (2-t-BP), 4-tert-butylphenol (4-t-BP), and others may also be leachable from plastic pipes.

**Keywords** Drinking water · Plastic pipe distribution system · Polyethylene pipe · Polypropylene pipe · Polybutylene pipe · Estrogenic compounds · Migration · Adverse effects

## Introduction

Since the 1980s, plastic piping has been increasingly utilized in drinking water distribution systems, in preference to other pipe materials such as cast iron, concrete, and copper. In Denmark, nearly all new drinking water pipes are made of polyethylene (PE), which comprised approximately 16% of the pipeline system in 2002. Meanwhile, 0.8–1.5% of the old pipeline system was replaced with PE pipes per year (Ryssel et al. 2015). According to the DK-VAND list of certified companies and products approved for drinking water in Denmark, pipes made of un-plasticized polyvinyl chloride (PVC) are also permitted for potable water system (DHI 2016).

According to Lucintel (2015), the global plastic pipe market has been forecast to expand further, with an annual growth rate of 6.8% from 2015 to 2020. The major drivers for this growing market are infrastructure development, rise in construction activities, replacement of aging pipelines made of traditional materials, and increasing population and urbanization. Potable water supply is expected to remain the largest market for plastic piping after wastewater supply application, as compared to alternative materials they are less expensive, lighter, and easier to install (Kelley et al. 2014; Lucintel 2015).

With the widespread application of plastic pipes in drinking water distribution systems, many substances which are potentially leachable from plastic pipes have been investigated (Brocca et al. 2002; Skjevraak et al. 2003; Loschner et al.

---

Responsible editor: Philippe Garrigues

✉ Ze-hua Liu  
zehualiu@scut.edu.cn

<sup>1</sup> School of Environment and Energy, South China University of Technology, Guangzhou, Guangdong 510006, China

<sup>2</sup> Key Lab Pollution Control & Ecosystem Restoration in Industry Cluster, Ministry of Education, Guangzhou, Guangdong 510006, China

<sup>3</sup> Guangdong Environmental Protection Key Laboratory of Solid Waste Treatment and Recycling, Guangzhou, Guangdong 510006, China

2011; Ryssel et al. 2015). To date, as many as 158 chemicals have been found to leach from plastic piping into drinking water (Whelton and Nguyen 2013). The migrating of chemicals from plastic pipes can result in a wide range of consequences, with the three main issues, increased total organic carbon (TOC) levels, increased odor problems in potable water, and potential public health risks, as outlined below.

TOC concentrations in drinking water are regulated in some countries as an indicator of drinking water quality. The maximum permitted TOC concentrations in Japanese and Chinese drinking water quality standards are 3 and 5 mg/L, respectively (Liu et al. 2016). Limits on TOC levels are mainly in place to minimize microbial growth in drinking water supplies, as the released TOC can be utilized by microorganisms as a source of energy and carbon, causing increased biofilm production in comparison to copper or stainless steel pipes (Van der Kooij et al. 2005).

Research by Kelley et al. (2014) revealed that the released TOC from cross-linked polyethylene (PEX) pipe was up to 6 mg/L in a simulated drinking water system, which exceeds the TOC limits for both Japanese and Chinese drinking water quality standards. There is though, some disagreement between studies as while Lund et al. (2011) found similar results to Kelley et al. (2014), other research has shown a much smaller effect on TOC levels by plastic piping (Heim and Dietrich 2007; Zhang et al. 2014).

The second main issue, increased odor problem in drinking water, is the result of plastics such as PEX piping producing a chemical/solvent odor in drinking water. A study by Durand and Dietrich (2007), identified 2-ethoxy-2-methylpropane (ETBE) as an important contributor to the odor. Kelley et al. (2014) monitored ETBE concentrations in a migration test and found ETBE to leach in a range of 3 to 179 µg/L during 1-year operation. As the odor threshold of ETBE is only 2 µg/L (Van Wezel et al., 2009), the abovementioned research in Kelly et al. (2014) indicates that drinking water odor issue due to ETBE migrating from plastic pipes may be a long-term problem.

In addition to ETBE, odorant migrating substances from plastics include methyl tert-butyl ether (MTBE), 1,3-dibutadiene and cyclohexadiene among others, may also exist in drinking water (Heim and Dietrich 2007; Lund et al. 2011; Kelley et al. 2014). In addition to the migrating compounds, the presence of their degradation byproducts as a result of biodegradation by biofilm microbes, may also be responsible for inducing strong odors over time (Skjevrak et al. 2005; Lund et al. 2011).

The third main problem with substances leaching from plastic pipelines is the potential human health risks these migrating compounds may cause. Compared to TOC and odor issues, there is less available research on the human health risk associated with migrating leachable chemicals. In one study, concentrations of ten migrating compounds were monitored, and their potential health risks were

evaluated based on their known toxicities, with the conclusion that the health risk from those migrating substances from new plastic piping, was low (Lund et al. 2011). The toxicity of two migrating compounds 2-t-BP and 2,4-d-t-BP to newborn rats, was studied by Hirata-Koizumi et al. (2005), establishing a no-observed-adverse-effect level (NOAEL) of 20 and 5 mg/kg/day, respectively. The high NOAELs of 2,4-d-t-BP appears to support the evidence that migrating compounds from plastic drinking water pipes have low potential health risks to humans. However, it should be noted that some migrating compounds such as 2-t-BP, 4-t-BP, and 2,4-d-t-BP are also reported to have estrogenic effects (Akahori et al. 2008; Tollefsen and Nilsen, 2008) and are reported to be endocrine disrupting compounds (EDCs).

In the last few decades, EDCs have been heavily researched due to their potential adverse effects on both wildlife and humans (Hu et al. 2009; Kidd et al. 2007; Liu et al. 2009a). A wide range of different substances belongs to the group of EDCs, including estrogens, androgens, phytoestrogens, progestins, as well as a number of industrial chemicals (Liu et al. 2011a, 2015a, 2015b). Due to the significant progress in EDC research, some of them have now been regulated within some drinking water quality standards.

Bisphenol A, diethyl phthalate, and di-butyl phthalate are three EDCs which are at present regulated in the Chinese drinking water quality standard, while 17β-estradiol (E2) and other four EDCs have been regulated in the latest Japanese drinking water quality standard (Liu et al. 2016). Despite this, little is known on their estrogenic potential, or the adverse effects of these estrogenic migrating compounds on human. Therefore, the main objective of this work was to evaluate the potential health risk of 2,4-d-t-BP from the viewpoint of endocrine disruption based on current existing literature, to further understand the effects of known and unknown estrogenic compounds, which are leachable from plastic drinking water pipes.

## Materials and methods

### Some basic information of 2,4-d-t-BP

2,4-d-t-BP is the most widely detected estrogenic compound leaching into drinking water from plastic pipes (Kelley et al. 2014; Loschner et al. 2011; Lund et al. 2011). The main source of 2,4-d-t-BP is from the degradation of phosphonite-based antioxidants such as antioxidant type 168 (Loschner et al. 2011), therefore, the presence of the antioxidants within plastic pipelines results in continual production of 2,4-d-t-BP, both within the pipe material or when antioxidant additives are released into

**Table 1** Basic physicochemical properties of 2,4-d-t-BP

Name	CAS	Molecular formula	Water solubility (mg/L)	LogKow	Boil point (°C)	Saturated pressure (mmHg)	Odor threshold (μg/L)	Estrogenic potencies	
								rtER	hER (α)
2,4-d-t-BP	96-76-4	C <sub>14</sub> H <sub>22</sub> O	120	4.97	263.5	0.00557	200 [1]	1.6e-5[2]	1.5e-5[3]

Information of the physicochemical properties is drawn from Scifinder, rtER: rainbow trout estrogen receptor; hER (α) in Table 1: human estrogen receptor α; [1] Tao and Zhang, 2010; [2] Tollefsen and Nilsen, 2008; [3] Akahori et al., 2008

drinking water. The basic physicochemical properties of 2,4-d-t-BP are outlined in Table 1.

### Potential health risk evaluation

In order to evaluate the potential health risks of 2,4-d-t-BP, firstly, we identified the existing levels of the target chemical in drinking water based on simulated migration test data available within scientific literature. The second step was to evaluate its potential risks, which was hindered significantly by the lack of direct investigation of the potential health risks associated with 2,4-d-t-BP as an estrogenic compound. Therefore, we evaluated comparable extensively studied estrogenic compounds, such as studies on potential health risks of 17α-ethynyl estradiol (EE2) and others at environmental relevant concentrations. However, the estrogenic activities of chemical compounds vary greatly, so for direct comparison

the concentrations of the estrogenic compounds were all expressed as estrogen equivalence (EEQ) as shown in the equation below (Liu et al. 2009b,2011b, Yuan et al., 2016).

$$EEQ = \sum EP_i \times c_i \quad (1)$$

Where EP and *c* denote the estrogenic potency of an individual estrogenic compound and the corresponding existing concentration, respectively. EP of estrogenic compound is often determined by different in vitro bioassays such as yeast estrogen screen (YES) and estrogen receptor (ER) competitive ligand binding assay, in which *E*<sub>2</sub> with the strongest estrogenic activity among natural estrogens is often selected as the standard estrogenic compound, i.e., the EP of *E*<sub>2</sub> is set as 1. When the estrogenic activity of one compound is stronger than that of *E*<sub>2</sub>, its EP would above 1, otherwise, the corresponding EP

**Table 2** Standard procedure studies assessing the migration levels of 2,4-d-t-BP concentration from plastic pipes into drinking water

No	Pipe type <sup>a</sup>	Pipe age <sup>b</sup> (d)	Detected concentration (μg/L)	EEQ (ngE2/L)	Oral exposure level (ngE2/d/P) <sup>d</sup>	Ref
1	PE	30	3.7 (23) <sup>c</sup>	0.06	0.12	[1]
2	PE-RT	30	2.8 (60)	0.04	0.08	[1]
3	PEXb	30	26 (23)	0.39	0.78	[1]
4	PEXc	30	10 (23); 31 (60)	0.15; 0.47	0.3; 0.94	[1]
5	PEXd	30	4.6 (60)	0.07	0.14	[1]
6	PP	30	6.1 (23); 13 (60)	0.09;0.2	0.18;0.4	[1]
7	PB	30	113 (23); >368 (60)	1.7; >5.5	3.4;>11	[1]
8	PEXd	3	0.5 (37)	0.01	0.02	[2]
9	HDPE	3	~5 (room)	~0.08	~0.16	[3]
10	PEXb	9	~0.68 (room)	~0.01	~0.02	[4]
11	PEXc	9	~2.2 (room)	~0.03	~0.06	[4]
12	PEXd	9	~2.9 (room)	~0.04	~0.08	[4]
13	PB	9	344.9 (room)	5.2	10.4	[4]
14	PEXd	150	0.37 (room)	0.01	0.02	[4]
15	PEXc	150	0.27 (room)	0.004	0.008	[4]

<sup>a</sup> PE-RT: polyethylene; PEXa: polyethylene cross-linked by peroxides; PEXb: polyethylene cross-linked by silanes; PEXc: polyethylene cross-linked by electron beam processing; PP: polypropylene; PB: polybutylene; HDPE: high density polyethylene

<sup>b</sup> Under simulated migration test

<sup>c</sup> Data in the parenthesis is the centigrade temperature

<sup>d</sup> Taken 2 L of drinking water is consumed every day, the EP value was based on hER (α); [1] Loschner et al. 2011; [2] Ryssel et al. 2015; [3] Skjjevrak et al. 2003; [4] Lund et al. 2011

**Table 3** Adverse effects of different EDCs on different kinds of model animals

No	Subject	Target chemical	Exposure level		Exposure time (d)	Main conclusions	Ref
			Concentration (ng/L)	EEQ <sup>a</sup> (ngEE2/L)			
1	Juvenile turbot ( <i>Psetta maxima</i> )	EE <sub>2</sub>	3.5	4.9	15	Reduction of both androgen production and plasma level in males was observed, which indicated that juvenile male turbot were susceptible to hormonal imbalance as a consequence of short-term exposure to environmentally relevant concentrations of EE <sub>2</sub> . [a]	
2	Japanese Medaka ( <i>Oryzias latipes</i> )	EE <sub>2</sub>	0.2–10	0.28–14	120–180 (posthatch to sexually mature)	1) At exposure level of 0.2–2 ng/L, mixed secondary sex characteristics in male fishes were observed, but mating behavior and reproductive success were normal; 2) when the exposure level increased to 10 ng/L, 16 among 19 males did not copulate, and reproductive success was very low; 3) female fish showed normal development and oogenesis even at exposure level of 10 ng/L, but they had poor reproductive success; 4) presence of oocytes in testicular tissue may not directly impact the reproductive capability of the male fish, but exposure of EDCs causing gonadal intersex was sufficient to reduce reproductive performance. [b]	
3	Adult zebrafish	EE <sub>2</sub>	10	14	17	1) Reduction in both male paternity and female maternity was observed; 2) reproductive success in male zebrafish was associated with the plasma concentration of 11-ketotestosterone and the corresponding concentration was suppressed in EE <sub>2</sub> -exposed males. [c]	
4	Embryonic zebrafish ( <i>Danio rerio</i> )	EE <sub>2</sub>	0.19–1	0.27–1.4	~240	1) No significant differences in transcription of vtg1 gene in zebrafish were observed in male fish at the exposure concentrations of 0.19 and 0.24 ng/L, while significantly elevated vtg1 gene transcription were observed at the exposure level of 1 ng/L; 2) a significant concentration-dependent increase in egg mortality between 8 and 24 h post-fertilization was observed at all the exposure concentrations of 0.19, 0.24, and 1 ng/L; 3) at two lowest exposure concentrations of 0.19 and 0.24 ng/L, EE <sub>2</sub> impacted late gastrulation and/or early organogenesis, whereas exposure to 1 ng/L disrupted development in the blastula phase. [d]	
5	Japanese medaka ( <i>Oryzias latipes</i> )	E <sub>2</sub>	29.3	29.3	25	At exposure concentration of 29.3 ng/L, 62.5% of male fishes had testis-ova gonad. [e]	
6	Adult zebrafish ( <i>Danio rerio</i> )	EE <sub>2</sub> , E <sub>2</sub> , E <sub>1</sub> , and other 8 EDCs	3–1608	20.7 (in total)	21	Vitellogenin mRNA and HIS in males confirmed both exposure regimes as physiological active. Meanwhile, potential candidates for estrogenic disturbance of steroidogenesis were identified (StAR, 17βhsd1, CYP19A1), which suggested that “non-classical effects” of estrogenic EDC in fish were mediated via transcription factor. [f]	
7	Frog ( <i>Xenopus tropicalis</i> )	EE <sub>2</sub>	1.8–180	2.52–252	32	EE <sub>2</sub> caused a concentration-dependent increase in the proportion of phenotypic females, and the percentages of females in control, 1.8, 18, and 180 ng/L of EE <sub>2</sub> were 32, 63, 90, and 100%, respectively. [g]	
8	Sydney rock oyster ( <i>Saccostrea glomerata</i> )	EE <sub>2</sub>	6.25–50	8.75–70	49	1) Vitellogenin was found to increase in a dose dependent manner with EE <sub>2</sub> exposure for males; [h]	

Table 3 (continued)

No	Subject	Target chemical	Exposure level		Exposure time (d)	Main conclusions	Ref
			Concentration (ng/L)	EEQ <sup>a</sup> (ngE2/L)			
9	Zebrafish ( <i>Danio rerio</i> )	E <sub>2</sub>	25	25	21	2) histological examination of gonads revealed a number of individuals exhibited intersex (ovotestis) at exposure level of 6.25 ng/L or more for 49 days. 1) Vitellogenin induction was observed in adult male fish under the E <sub>2</sub> exposure concentrations between 5 and 25 ng/L; 2) a modification of the secondary sexual characteristics in adult male fish at exposure concentration of 25 ng/L was observed.	[i]
10	Male fathead minnows	4-nonylphenol	6100	1.84	14	1) Exposure resulted in an induction of plasma vitellogenin in males within 14 days; 2) reproductive competence of the exposed males was decreased when compared to the controlled males.	[j]
11	Salmon smolts	4-nonylphenol	2000	0.60	21	1) Exposure of 4-nonylphenol would result in delayed mortality rates; 2) One year after exposure, yolk-sac larvae decreased gill sodium-potassium-activated adenosine triphosphatase activity and seawater tolerance during smolt development; 3) exposed fish had 20% lower plasma insulin-like growth factor I levels and 35% lower plasma triiodothyronine.	[k]
12	Zebrafish ( <i>Danio rerio</i> )	EE <sub>2</sub>	5	7	Life-long exposure	1) Life-long exposure caused a 56% reduction in fecundity and complete population failure with no fertilization; 2) infertility was due to disturbed sexual differentiation, with males having no functional testes and either undifferentiated or intersex gonads.	[l]
13	Mussel ( <i>Mytilus edulis</i> )	Bisphenol A	50,000	6.3	21	1) It induced the expression of phospho-proteins in females and spawning in both sexes; 2) Severe damaging effects on ovarian follicles and oocytes were observed in female mussels.	[m]
14	Zebrafish	Zearalenone	100–1000	2.83–28.3	21	1) At exposure concentration of 1000 ng/L, the relative spawning frequency was reduced to 38.9%; 2) relatively fecundity at 100 and 1000 ng/L was reduced to 74.2 and 43.8%, respectively; 3) a 4–4-fold induction of plasma vitellogenins was observed in male zebrafish at 1000 ng/L.	[n]

<sup>a</sup> EP values of EE<sub>2</sub>, 4-nonylphenol, bisphenol A and zearalenone were  $1.4$ ,  $3.01 \times 10^{-4}$ ,  $1.26 \times 10^{-4}$ , and  $2.83 \times 10^{-2}$ , respectively, which were derived from Liu et al. (2009c, 2010); [a] Labadie and Budzinski, 2006; [b] Balch et al. 2004; [c] Coe et al. 2008; [d] Soares et al. 2009; [e] Kang et al. 2002; [f] Urbazka et al. 2012; [g] Pettersson and Berg, 2007; [h] Andrew et al. 2010; [i] Brion et al. 2004; [j] Schoenfuss et al. 2008; [k] Lerner et al. 2007; [l] Nash et al. 2004; [m] Arab et al., 2006; [n] Schwartz et al., 2010

would be below 1. Therefore, the unit of EEQ is normally represented as ngE<sub>2</sub>/L.

## Results and discussions

### Migration levels

Polymer plastic materials may include numerous residual monomers, oligomers, or other substances and additives (such as antioxidants) for protection of pipe material during its production and use. Many of these substances, as well as their degradation products and impurities have the potential to migrate into the drinking water (Loschner et al. 2011). In order to check potential migrating chemicals, some standard procedures such as EN-1420-1, prEN-15768, and NSF1 standard 61 have been applied, to simulate the migration levels of substances from plastic pipes into drinking water (Loschner et al. 2011; Lund et al. 2011; Kelley et al. 2014). In these tests, a pre-flushed new plastic pipe is filled with ultrapure water, allowed to stagnate for 72 h and then emptied, with the effluent collected and prepared for the analysis of the migrating compounds. To evaluate the long-term period migration, the above migration test is continued with repeated ultrapure water testing every 72 h. The results of these studies are summarized in Table 2.

As shown in Table 2, several kinds of plastic pipes have been utilized in drinking water distribution systems. PB pipe has been shown to release the highest concentrations of 2,4-d-t-BP into drinking water among all plastic pipe materials. Studies have shown that the release of 2,4-d-t-BP from PEX pipes is more complex. Loschner et al.

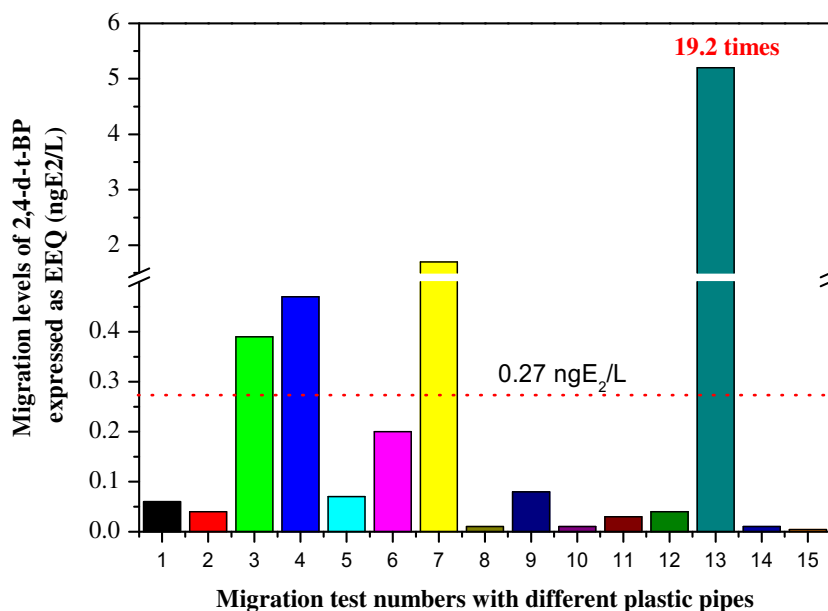
(2011), found three kinds of PEX pipes to release between 4.6–26 μg/L of 2,4-d-t-BP into drinking water after 1 month of pipe usage.

One potential reason for the long-term release of 2,4-d-t-BP at relatively high levels from PEX pipes, might be that the compound is non-biodegradable to the microbial communities that form the biofilms within PEX plastic pipes (Ryssel et al. 2015). However, another study found that the same three kinds of new PEX pipes, after 9 days of usage released no more than 2.9 μg/L of 2,4-d-t-BP (Lund et al. 2011). As the temperature between the two above simulated tests were similar, these results seem to suggest that PEX pipes made of the same material but by different manufacturing brands, may leach varying amounts of 2,4-d-t-BP into drinking water. This variation should be investigated further as it may also apply to other plastic pipes.

### Potential health risk evaluation

It is of some concern that 2,4-d-t-BP is widely used in food-related plastic products as an antioxidant. Since 1967, the US Food and Drug Administration have permitted its usage in packaging for irradiated foods (Paquette 2004) and since the 2011 ban on the production of polycarbonate baby bottles within the EU, due to the toxic effect of bisphenol A, 2,4-d-t-BP has been detected in baby bottles made of the alternative approved materials, despite the fact that it has not been safety assessed under the European Union Regulations (10/2011) (Onghena et al. 2014). Due to the lack of direct investigations, the potential health risks of 2,4-d-t-BP are evaluated based on the EEQ value.

**Fig. 1** The migration levels of 2,4-d-t-BP from different plastic pipes expressed as EEQ under simulated migration test (numbers in the horizontal axis agree with the numbers of the migration tests in Table 2)



There have been various studies on the adverse effects of EDCs on fish and other animal model species, at environmentally relevant concentrations. These studies have been summarized, as shown in Table 3. EE<sub>2</sub> is at present the most widely studied EDC and the adverse effects mainly include vitellogenin induction, intersex induction (the development of ovotestis), infertility, and disruption to mating behaviors, in addition to mortality and other effects. It is also significant that different animal models or different developmental stages show significant variation in their responses to EDC exposure. However, trace EEQ concentrations of less than 10 ngE<sub>2</sub>/L, resulted in adverse effects in ten out of 14 (71.4%) of the summarized studies in Table 3, including effects on seven different animal models and developmental stages, including juvenile turbot, Japanese medaka, zebrafish, frog; Sydney rock oysters, fathead minnows, and salmon smolts. Soares et al. (2009) reported that at EEQ exposure levels as low as 0.27 ngE<sub>2</sub>/L, egg mortality and other adverse effects could be seen in late gastrulation and/or early organogenesis in zebrafish. Therefore, the EEQ levels of 2,4-d-t-BP leached from the plastic pipes in four out of the 15 simulated drinking water tests (Fig. 1), would comparatively result in a higher level of adverse effects than seen in the study by Soares et al. (2009). This is especially the case for PB pipes where the maximal migrating EEQ level of 2,4-d-t-BP at room temperature was 5.2 ng E<sub>2</sub>/L, over 19 times the EEQ level that caused adverse effects on the zebrafish model in the study by Soares et al. (2009). Although PB pipes are not as widely used as PEX pipes in drinking water distribution system, in some applications PB pipes are suggested as replacement for PEXs (Lund et al. 2011). This is of concern as the human consumption of 2 L drinking water per day could result in a daily 2,4-d-t-BP oral exposure of up to 10.4 ngE<sub>2</sub>/d/P.

It is generally assumed that potable drinking water, such as those simulated in Table 2, are safe for human consumption. However, the EEQ level of 2,4-d-t-BP in some drinking water supplies, raise serious questions about the potential health effects of long-term consumption of such drinking water transported with plastic pipes.

It must be considered that EDCs with the same EEQ levels may still induce very different adverse effects on animal models, thus the reported results of the adverse effects summarized in Table 3 may not necessarily reflect the same adverse effects caused by 2,4-d-t-BP in drinking water. In most cases, the migrating levels of 2,4-d-t-BP in PE and PEX pipes are relative low, and only 15.3% of the tested pipes leached levels of 2,4-d-t-BP that may cause adverse health effects. However, due to the tight regulations for drinking water, further investigation of this contaminant should be undertaken and under the current strict requirements on drinking water quality; the latent risks of 2,4-d-t-BP as an estrogenic compound must be better understood and regulated. Moreover, 2,4-d-t-BP is not the only estrogenic compound leachable from the plastic pipes, other migrating compounds such as 2-t-BP and 4-t-BP also showed

estrogenic activities (Akahori et al. 2008), suggesting that this problem is more complex than removing one harmful substance from our water supplies. Given our limited knowledge on the migrating compounds, more attentions should be paid to the safety of drinking water from the viewpoint of its estrogenicity.

## Conclusions

The migrating levels of 2,4-d-t-BP into drinking water from different kinds of plastic pipes under simulated migration tests were summarized. Generally, the PB pipes released much higher levels of 2,4-d-t-BP than PE and PP pipes, and plastic pipes made of the same materials but by different manufacturing brands may release different amounts of 2,4-d-t-BP. To evaluate the latent risk effects of 2,4-d-t-BP, the migrating levels from different plastic pipes were converted into EEQ levels. Although the EEQ levels of 2,4-d-t-BP migrating from pipes in most occasions were relative low. However, in a few cases, the released levels of 2,4-d-t-BP may cause potential adverse health effects on humans as the maximal migrating levels were over 19 times higher than the level that could pose adverse effects to different model animals. In addition, it should be pointed out that the continual release of 2,4-d-t-BP from plastic pipes may last for 1 year or longer, which suggest that the potential adverse influence may not be a short-term problem. To the best of our knowledge, this is the first report of estrogenic levels of 2,4-d-t-BP in drinking water, a readily leachable compound from plastic pipes that widely used in drinking water distribution systems.

**Acknowledgement** This work was financially supported by The Key Program of National Natural Science of China (No. 41330639); the Program for National Natural Science Foundation of China (No. 21107025; No.21577040; U1501234); Special funds for public welfare research and capacity building in Guangdong Province (B2153210); Science and Technology Program of Guangzhou, China (No. 201510010162); the Fundamental Research Funds for the Central Universities (2014ZM0073), as well as the funding for water odor research (D8144320).

## References

- Akahori Y, Nakai M, Yamasaki K, Takatsuki M, Shimohigashi Y, Ohtaki M (2008) Relationship between the results of in vitro receptor binding assay to human estrogen receptor  $\alpha$  and in vivo uterotrophic assay: comparative study with 65 selected chemicals. *Toxicol in Vitro* 22:225–231
- Andrew MN, O'Connor WA, Dunstan RH, MacFarlane GR (2010) Exposure to 17 $\alpha$ -ethynylestradiol causes dose and temporally dependent changes in intersex, females and vitellogenin production in the Sydney rock oyster. *Ecotoxicology* 19:1440–1451
- Arab N, Lemaire-Gony S, Unruh E, Hansen PD, Larsen BK, Andersen OK, Narbonne JF (2006) Preliminary study of responses in mussel (*Mytilus edulis*) exposed to bisphenol A, diallyl phthalate and tetrabromodi phenyl ether. *Aquat Toxicol* 78S:S86–S92

- Balch GC, Mackenzie CA, Metcalfe CD (2004) Alterations to gonadal development and reproductive success in Japanese medaka (*Oryzias latipes*) exposed to 17 $\alpha$ -ethynylestradiol. *Environ Toxicol Chem* 23:782–791
- Brion F, Tyler CR, Palazzi X, Laillet B, Porcher JM, Garric J, Flammarion P (2004) Impacts of 17 $\beta$ -estradiol, including environmentally relevant concentrations, on reproduction after exposure during embryonal-, juvenile- and adult-life stages in zebrafish (*Danio rerio*). *Aquat Toxicol* 68:193–217
- Brocca D, Arvin E, Mosbaek H (2002) Identification of organic compounds migrating from polyethylene pipelines into drinking water. *Water Res* 36:3675–3680
- Coe TS, Hamilton PB, Hodgson D, Paul GC, Stevens JR, Sumner K, Tyler CR (2008) An environmental estrogen alters reproductive hierarchies, disrupting sexual selection in group-spawning fish. *Environmental science & technology* 42:5020–5025
- DHI (2016) [www.dancert.dk/english/plastic-pipe-systems-for-drinking-water-supply-in-denmark/35790](http://www.dancert.dk/english/plastic-pipe-systems-for-drinking-water-supply-in-denmark/35790). Access in 21 Apr 2016
- Durand ML, Dietrich AM (2007) Contribution of silane cross-linked PEX pipe to chemical/solvent odours in drinking water. *Wat Sci Technol* 55:153–160
- Heim TH, Dietrich AM (2007) Sensory aspects and water quality impacts of chlorinated and chloraminated drinking water in contact with HDPE and PVC pipe. *Water Res* 41:757–764
- Hirata-Koizumi M, Hamamura M, Furukawa H, Fukuda N, Ito Y, Wako Y, Yamashita K, Takahashi M, Kamata E, Ema M, Hashigawa R (2005) Elevated susceptibility of newborn as compared with young rats to 2-tert-butylphenol and 2,4-di-tert-butylphenol toxicity. *Congenital anomalies* 45:146–153
- Hu JY, Zhang ZB, Wei QW, Zhen HJ, Zhao YB, Peng H, Wan Y, Giesy JP, Li LX, Zhang B (2009) Malformation of the endangered sturgeon, *Acipenser sinensis*, and its causal agent. *Proc Natl Acad Sci U S A* 106:9339–9344
- Kang JJ, Yokota H, Oshima Y, Tsuruda Y, Yamaguchi T, Maeda M, Imada N, Tadokoro H, Honjo T (2002) Effect of 17 $\beta$ -estradiol on the reproduction of Japanese medaka (*Oryzias latipes*). *Chemosphere* 47:71–80
- Kelley KM, Stenson AC, Dey R, Whelton AJ (2014) Release of drinking water contaminants and odor impacts caused by green building cross-linked polyethylene (PEX) plumbing systems. *Water Res* 67:19–32
- Kidd KA, Blanchfield PJ, Mills KH, Palace VP, Evans RE, Lazorchak JM, Flick RW (2007) Collapse of a fish population after exposure to a synthetic estrogen. *Proc Natl Acad Sci U S A* 104:8897–8901
- Labadie P, Budzinski H (2006) Alteration of steroid hormone profile in juvenile turbot (*Psetta maxima*) as a consequence of short-term exposure to 17 $\alpha$ -ethynyl estradiol. *Chemosphere* 64:1274–1286
- Lerner DT, Bjornsson BT, McCormick SD (2007) Larval exposure to 4-nonylphenol and 17 $\beta$ -estradiol affects physiological and behavioral development of seawater adaption in Atlantic salmon smolts. *Environmental science & technology* 41:4479–4485
- Liu ZH, Kanjo Y, Mizutani S (2009a) Urinary excretion rates of natural estrogens and androgens from humans, and their occurrence and fate in the environment: a review. *Sci Total Environ* 407:4975–4985
- Liu ZH, Ito M, Kanjo Y, Yamamoto A (2009b) Profile and removal of endocrine disrupting chemicals by using an ER/AR competitive ligand binding assay and chemical analysis. *J Environ Sci* 21:900–906
- Liu ZH, Kanjo Y, Mizutani S (2009c) Removal mechanisms of endocrine disrupting compounds (EDCs) in wastewater treatment-physical means, biodegradation, and chemical advanced oxidation: a review. *Sci Total Environ* 407:731–748
- Liu ZH, Kanjo Y, Mizutani S (2010) A review of phytoestrogens: their occurrence and fate in the environment. *Water Res* 44:567–577
- Liu ZH, Ogejo JA, Pruden A, Knowlton KF (2011b) Occurrence, fate and removal of synthetic oral contraceptives (SOCs) in the natural environment: a review. *Sci Total Environ* 409:5149–5161
- Liu ZH, Kanjo Y, Mizutani S (2011a) Removal of natural free estrogens and their conjugates in a municipal wastewater treatment plant. *Clean-soil air water* 39:128–135
- Liu ZH, Lu GN, Yin H, Dang Z, Littler H, Liu Y (2015a) Sample-preparation methods for direct and indirect analysis of natural estrogens. *TrAC - Trends in Analytical Chemistry* 64:149–164
- Liu ZH, Lu GN, Yin H, Dang Z, Rittmann B (2015b) Removal of natural estrogens and their conjugates in municipal wastewater treatment plants: a critical review. *Environ Sci Technol* 49:5288–5300
- Liu ZH, She PY, Wei XN, Deng L, Dang Z (2016) New drinking water standards in Japan and its discussions. *China Water & Wastewater* 32:8–10 (in Chinese)
- Loschner D, Rapp T, Schlosser FU, Schuster R, Stottmeister E, Zander S (2011) Experience with the application of the draft European Standard prEN 15768 to the identification of leachable organic substances from materials in contact with drinking water by GC-MS. *Anal Methods* 3:2547–2556
- Lucintel (2015) Growth opportunities in global plastic pipe market 2015–2020: trend, forecast, and market analysis. [www.lucintel.com/plastic\\_pipe\\_market\\_2020.aspx](http://www.lucintel.com/plastic_pipe_market_2020.aspx). Accessed in 21 Apr 2016
- Lund V, Anderson-Glenna M, Skjevrak I, Steffensen IL (2011) Long-term study of migration of volatile organic compounds from cross-linked polyethylene (PEX) pipes and effects on drinking water quality. *Journal of water health* 9:483–497
- Nash JP, Kime DE, Van der Ven LTM, Wester PM, Brion F, Macck G, Stahlschmidt-Allner P, Tyler CR (2004) Long-term exposure to environmental concentrations of the pharmaceutical ethynylestradiol causes reproductive failure in fish. *Environmental health perspective* 112:1725–1733
- Ongghena M, van Hoeck E, Vervliet P, Scippo ML, Simon C, van Loco J, Covaci A (2014) Development and application of a non-targeted extraction method for the analysis of migrating compounds from plastic baby bottles by GC-MS. *Food additives and contaminants part A-chemistry analysis control exposure & risk assessment* 31:2090–2102
- Paquette KE (2004) Irradiation of prepackaged food: evolution of the US food and drug administration's regulation of the packaging materials. *ACS Symp Ser* 85:182–202
- Pettersson I, Berg C (2007) Environmentally relevant concentrations of ethynylestradiol cause female-biased sex ratios in *Xenopus tropicalis* and *Rana temporaria*. *Environ Toxicol Chem* 26:1005–1009
- Ryssel ST, Arvin E, Lutzhoft HCH, Olsson ME, Procazkova Z, Albrechtsen HJ (2015) Degradation of specific aromatic compounds migrating from PEX pipes into drinking water. *Water Res* 81:269–278
- Schoenfuss HL, Bartell SE, Bistodeau TB, Cediell RA, Grove KJ, Zinte KL, Lee KE, Barber LB (2008) Impairment of the reproductive potential of male fathead minnows by environmentally relevant exposures to 4-nonylphenol. *Aquat Toxicol* 86:91–98
- Schwartz P, Thorpe KL, Bucheli TD, Wettstein FE, Burkhardt-Holm P (2010) Short-term exposure to the environmentally relevant estrogenic mycotoxin zearlenone impairs reproduction in fish. *Sci Total Environ* 409:326–333
- Skjevrak I, Due A, Gjerstad KO, Herikstad H (2003) Volatile organic components migrating from plastic pipes (HDPE, PEX, and PVC) into drinking water. *Water Res* 37:1912–1920
- Skjevrak I, Lund V, Ormerod K, Herikstad H (2005) Volatile organic compounds in natural biofilm in polyethylene pipes supplied with lake water and treated water from the distribution network. *Water Res* 39:4133–4141
- Soares J, Coimbra AM, Reic-Henriques MA, Monteiro NM, Vieira MN, Oliveira JMA, Guedes-Dias P, Fontainhas-Fernandes A, Parra SS, Carvalho AP, Castro FC, Santos MM (2009) Disruption of zebrafish (*Danio rerio*) embryonic development after full life-cycle parental exposure to low levels of ethynylestradiol. *Aquat Toxicol* 95:330–338
- Tao Y, Zhang L (2010) Intensity prediction of typical aroma characters of cabernet sauvignon wine in Changli County (China). *LWT Food Sci Technol* 43:1550–1556



- Tollefsen KE, Nilsen AJ (2008) Binding of alkylphenols and alkylated non-phenolics to rainbow trout (*Oncorhynchus mykiss*) hepatic estrogen receptors. *Ecotoxicol Environ Saf* 69:163–172
- Urbatzka R, Rocha E, Reis B, Cruzeiro C, Monteiro RAF, Rocha MJ (2012) Effects of ethinylestradiol and of an environmentally relevant mixture of xenoestrogens on steroidogenic gene expression and specific transcription factor in zebrafish. *Environ Pollut* 164:28–35
- Van der Kooij D, Veenendaal HR, Scheffer WJH (2005) Biofilm formation and multiplication of legionella in a model warm water system with pipes of copper, stainless steel and cross-linked polyethylene. *Water Res* 39:2789–2798
- Van Wezel A, Puijker L, Vink C, Versteegh A, de Voogt P (2009) Odour and flavor thresholds of gasoline additives (MTBE, ETBE and TAME) and their occurrence in Dutch drinking water collection areas. *Chemosphere* 76:672–676
- Whelton AJ, Nguyen T (2013) Contaminant migration from polymeric pipes used in buried potable water distribution systems: a review. *Crit Rev Sci Technol* 43:679–751
- Yuan SF, Liu ZH, Lian HX, Yang CT, Lin Q, Yin H, Dang Z (2016) Simultaneous determination of estrogenic odorant alkylphenols, chlorophenols, and their derivatives in water using online headspace solid phase microextraction coupled with gas chromatography-mass spectrometry. *Environ Sci Pollut Res* 23:19116–19125
- Zhang L, Liu SM, Liu WJ (2014) Investigation of organic matter migrating from polymeric pipes into drinking water under different flow manners. *Environmental science process & impacts* 116:280–290