

The Prospective Toxic Effects of Some Heavy Metals Overload in Surface Drinking Water of Dakahlia Governorate, Egypt

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Abstract

Background: The concentration of heavy metals in drinking water is very important.

Objectives: To evaluate the chemistry of some heavy metals in surface drinking water of Dakahlia Governorate, Egypt.

Methods: 51 surface drinking water samples were collected from the main surface water stations and compact units in October 2009 and analyzed chemically. 26 water samples were analyzed by atomic absorption spectrophotometer for iron, manganese, lead, nickel, chromium, zinc, copper, cobalt, aluminum, and cadmium concentrations.

Results: Aluminum concentration was slightly high in water sample of Bosat network. Cadmium concentration in samples of up-streams of shark and Mit-Khamis stations, networks of Mit-Antar, Demera, Bosat, Bilqas, El-satamony, El-Gamalia, Mit-asim and Bilqas station exceeded the permissible limits of Egyptian Ministry of Health (EMH, 2007) and World Health Organization (WHO, 2008). The nickel concentration in the network samples taken from Shoha, Bosat and El-Gamalia as well as up-stream of Bosat station exceeded the permissible limits. Also, lead concentrations of the network samples of Shoha, Mit-Antar, Demera and Nabaru exceeded the permissible limits.

Conclusion: Regular chemical analysis of surface drinking water is required. Since these heavy metals are most likely originate from steel, plastics and batteries industries working in the region, we believe that activities of these industries must be stopped or at least limited in urban zones.

Keywords: Surface water; Aluminum; Cadmium; nickel; Metals, Heavy; Elements

Introduction

Dakahlia Governorate is a region in North of Egypt. The surface water in Dakahlia Governorate is mainly from the River Nile flowing in Damietta branch and its distributaries irrigation canals. Abd el-Daiem and Ramsussen reported that at Mansoura, the surface water level in the Nile branch is +1.8 m, while it

is about +4 m in the nearby water wells, where the river is recharged by groundwater and becomes effluent stream in that sector and the Nile branch down stream of Mansoura behaves as a drain.¹ Rainwater collects impurities while passing through the air. Streams and rivers collect impurities from surface run-off and through the discharge of sewage, industrial and agriculture effluents. These are carried to the

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Table 1: Chemical analysis in ppm (mg/L) of water samples taken from the main surface drinking water stations

| No | District/sample name | pH | EC $\mu\text{m/cm}$ at 25 °C | TDS ppm | TH ppm | Cl ₂ ppm | Cl ⁻ ppm | SO ₄ ²⁻ ppm | Alkal. ppm | Al ³⁺ ppm |
|-----|--|----------------|---------------------------------|-------------|------------|------------------------|------------------------|--------------------------------------|---------------|-------------------------|
| | Permissible limit of WHO (2008) | 6.5–8 | | 1000 | 500 | 5 | 250 | 250 | | 0.1 |
| | Permissible limit of EMH (2007) | 6.5–8.5 | | 1000 | 500 | 5 | 250 | 250 | | 0.2 |
| | El-Mansoura district | | | | | | | | | |
| I | Up-stream of shark station | 8.18 | 414 | 269.1 | 136 | — | 22.7 | 67.31 | 155 | — |
| | Main station | 7.75 | 444 | 288.6 | 144 | 0.66 | 35.45 | 75 | 150 | 0.03 |
| | Network | 7.85 | 450 | 292.5 | 140 | 0.52 | 34.75 | 81.73 | 140 | 0.04 |
| | Up-stream of Mit-khamis station | 8.05 | 430 | 279 | 145 | — | 25.53 | 66 | 160 | — |
| | Main station | 7.69 | 443 | 288 | 136 | 1.56 | 34.73 | 99 | 145 | 0.08 |
| | Network | 7.6 | 446 | 290 | 137 | 1.5 | 34.03 | 80 | 140 | 0.03 |
| | Talkha district | | | | | | | | | |
| II | Up-stream of Talkha station | 8.0 | 427 | 277.5 | 137 | — | 24.11 | 34 | 151 | — |
| | Main station | 7.68 | 425 | 276.3 | 144 | 1.95 | 35.45 | 60 | 146 | 0.10 |
| | Network | 7.75 | 436 | 283.4 | 136 | 1.3 | 34.03 | 46 | 142 | 0.05 |
| | Dekernis district | | | | | | | | | |
| III | Up-stream of Mit-Faris station | 7.99 | 426 | 276.9 | 140 | — | 29.8 | 43.3 | 160 | — |
| | Main station | 7.65 | 428 | 278.2 | 132 | 0.53 | 32.6 | 90 | 135 | 0.09 |
| | Network | 7.72 | 461 | 269.6 | 142 | 0.33 | 34.03 | 78.85 | 150 | 0.08 |
| | Sherbin district | | | | | | | | | |
| IV | Up-stream of Sherbin station | 8.05 | 440 | 286 | 144 | — | 31.2 | 39 | 170 | — |
| | Main station | 7.74 | 438 | 284.7 | 137 | 1.5 | 34.03 | 56 | 155 | 0.10 |
| | Network | 7.41 | 433 | 281.5 | 141 | 1.35 | 33.33 | 44 | 152 | 0.05 |
| | Up-stream of Bosat station | 8.23 | 448 | 291.2 | 144 | — | 35.45 | 76.93 | 165 | — |
| | Main station | 7.73 | 435 | 282.2 | 128 | 0.6 | 36.9 | 75 | 150 | 0.10 |
| | Network | 7.8 | 460 | 299 | 140 | 0.33 | 38.28 | 60.57 | 175 | 0.11 |
| | Bilqas district | | | | | | | | | |
| V | Up-stream of Bilqas station | 8.05 | 462 | 300.3 | 142 | — | 34.03 | 33.46 | 160 | — |
| | Main station | 7.76 | 477 | 310.1 | 144 | 0.64 | 35.45 | 96.15 | 155 | 0.05 |
| | Network | 7.81 | 475 | 308.7 | 140 | 0.41 | 38.3 | 86.54 | 150 | 0.08 |

Continued

Table 1: Chemical analysis in ppm (mg/L) of water samples taken from the main surface drinking water stations

| No | District/sample name | pH | EC $\mu\text{m/cm}$ at 25 °C | TDS ppm | TH ppm | Cl ₂ ppm | Cl ⁻ ppm | SO ₄ ²⁻ ppm | Alkal. ppm | Al ³⁺ ppm |
|--|-------------------------------------|----------------|---------------------------------|-------------|------------|------------------------|------------------------|--------------------------------------|---------------|-------------------------|
| Permissible limit of WHO (2008) | | 6.5–8 | | 1000 | 500 | 5 | 250 | 250 | | 0.1 |
| Permissible limit of EMH (2007) | | 6.5–8.5 | | 1000 | 500 | 5 | 250 | 250 | | 0.2 |
| El-Gamalia district | | | | | | | | | | |
| VI | Up-stream of El-Gamalia station | 8.14 | 428 | 278.2 | 134 | — | 31.2 | 73.07 | 150 | — |
| | Main station | 7.73 | 435 | 282.7 | 134 | 0.57 | 35.45 | 79.81 | 125 | 0.05 |
| | Network | 7.62 | 438 | 284.7 | 130 | 0.36 | 32.6 | 86.5 | 150 | 0.06 |
| El-Sinbillawin district | | | | | | | | | | |
| VII | Up-stream of El-Sinbillawin station | 7.15 | 420 | 270 | 134 | — | 30.1 | 71 | 138 | — |
| | Main station | 7.35 | 430 | 275 | 135 | 0.76 | 36.1 | 76 | 120 | 0.07 |
| | Network | 7.12 | 434 | 280 | 130 | 0.32 | 32.1 | 85 | 144 | 0.04 |
| El-Manzala district | | | | | | | | | | |
| VIII | Up-stream of El-Manzala station | 8.11 | 377 | 245 | 140 | — | 21 | 32 | 145 | 0.01 |
| | Main station | 7.45 | 390 | 253 | 124 | 0.08 | 20 | 35 | 130 | 0.05 |
| | Network | 7.75 | 393 | 255 | 136 | 0.18 | 33 | 53 | 151 | 0.06 |

ppm: Part per million = mg/L, EC: Electrical conductivity $\mu\text{m/cm}$: $\mu\text{mohs/cm}$, TDS: Total dissolved salts, TH: Total hardness, Cl₂: Chlorine, Cl⁻: Chloride, SO₄²⁻: Sulfate, Alkal: Alkalinity

rivers, lakes or reservoirs that supply our drinking water.² Cadmium (Cd), copper (Cu), cobalt (Co), chromium (Cr), manganese (Mn), nickel (Ni), lead (Pb), and zinc (Zn) are toxicogenic consistently found as contaminants in human drinking water supplies in many areas around the world.³

We conducted this study to evaluate the chemistry of surface drinking water with special emphasis on the concentrations of some heavy metals; iron (Fe), Mn, Pb, Ni, Cr, Zn, Cu, Co and Cd and their toxic impacts on human health.

Materials and Methods

Fifty-one surface drinking water samples were taken from the main station (n=30) and compact units (n=21) in Dakahlia Governorate in October 2009. All samples were collected in 2-L polyethylene bottles for the chemical analysis. The parameters, pH, electrical conductivity (EC) and total dissolved salts (TDS) were measured in the field.

The pH was measured by a pH meter (HI 8424 microcomputer, Hanna instru-

ments). TDS and EC were measured using conductivity/TDS-meter (ESD, Engineered Systems & Design). The 51 water samples were subjected to chemical analysis according to the procedure described in the standard method for the examination of water and wastewater.⁴ Twenty-six water samples were subjected for measuring levels of Fe, Mn, Pb, Ni, Cr, Zn, Cu, Co and Cd by atomic absorption spectrophotometer (AAS) (Buck Scientific Company,

USA). Measurements of each metal level were carried out in duplicate with the suitable parameters (wavelength, concentration ranges and detection limit).

Results

Tables 1 and 2 show that the measured pH values were within the permissible limits of Egyptian Ministry Health (EMH) permissible limits,⁵ however they exceed

Table 2: Results of chemical analysis in ppm of the compact units

| No | District/sample name | pH | EC $\mu\text{m/cm}$ at 25 °C | TDS ppm | TH ppm | Cl ₂ ppm | Cl ⁻ ppm | SO ₄ ²⁻ ppm | Alkal. ppm | Al ³⁺ ppm |
|----------------------|----------------------|------|------------------------------|---------|--------|---------------------|---------------------|-----------------------------------|------------|----------------------|
| El-Mansoura district | | | | | | | | | | |
| I | Awish el-hagar | 7.53 | 486 | 315.9 | 150 | 0.4 | 35 | 43 | 148 | 0.03 |
| | Mahalet damana | 7.82 | 420 | 273 | 126 | 0.28 | 32 | 52 | 136 | 0.02 |
| | Shoha | 7.7 | 371 | 241 | 130 | 0.12 | 30 | 44 | 131 | 0.02 |
| | El-Rydanian | 7.77 | 410 | 266 | 130 | 0.21 | 40 | 56 | 143 | 0.02 |
| Talkha district | | | | | | | | | | |
| II | Mit-antar | 7.46 | 445 | 298.5 | 145 | 1.5 | 34.03 | 65 | 156 | 0.06 |
| | Mit- elkorama | 7.63 | 486 | 315.9 | 150 | 0.5 | 31 | 67 | 140 | 0.01 |
| | Demera | 7.74 | 468 | 304.2 | 130 | 0.45 | 35 | 46 | 120 | 0.03 |
| Nabaru district | | | | | | | | | | |
| III | Nabaru | 7.67 | 480 | 312 | 140 | 0.71 | 50 | 60 | 145 | 0.06 |
| Sherbin district | | | | | | | | | | |
| IV | Ras el-khalig | 7.72 | 498 | 323.7 | 140 | 0.6 | 46 | 57 | 133 | 0.04 |
| Bilqas district | | | | | | | | | | |
| V | El-satamony | 7.99 | 385 | 256 | 135 | 0.4 | 30 | 36 | 141 | 0.04 |
| | Basindela | 7.46 | 410 | 266.5 | 135 | 0.15 | 30 | 62 | 130 | 0.01 |
| Aga district | | | | | | | | | | |
| VI | Nawasa | 7.53 | 398 | 259 | 130 | 0.12 | 21 | 30 | 140 | 0.01 |

Continued

Table 2: Results of chemical analysis in ppm of the compact units

| No | District/sample name | pH | EC $\mu\text{m/cm}$ at 25 °C | TDS ppm | TH ppm | Cl ₂ ppm | Cl ⁻ ppm | SO ₄ ²⁻ ppm | Alkal. ppm | Al ³⁺ ppm |
|-------------------------|----------------------|------|------------------------------|---------|--------|---------------------|---------------------|-----------------------------------|------------|----------------------|
| El-Manzala district | | | | | | | | | | |
| VII | El-aziza | 7.79 | 379 | 246 | 140 | 0.1 | 28 | 45 | 145 | — |
| | El-Mawaged | 7.44 | 375 | 243 | 132 | 0.48 | 30 | 45 | 140 | 0.02 |
| Mit-Salsil district | | | | | | | | | | |
| VIII | Mit-Salsil | 7.9 | 447 | 290.5 | 140 | 0.66 | 34 | 57 | 180 | 0.04 |
| El-Sinbillawin district | | | | | | | | | | |
| IX | Barqin | 7.73 | 359 | 233 | 135 | 0.4 | 30 | 45 | 140 | 0.03 |
| Temy el-amdid district | | | | | | | | | | |
| X | Temy el-amdid | 7.76 | 454 | 295.1 | 160 | 0.43 | 38 | 51 | 150 | 0.01 |
| | Abo-dawoud | 8.01 | 432 | 280.8 | 146 | 0.3 | 36 | 42 | 150 | 0.01 |
| Minyet el-nasr district | | | | | | | | | | |
| XI | Minyet el-nasr | 7.5 | 431 | 281.2 | 130 | 0.8 | 30 | 47 | 136 | 0.06 |
| | Mit-asim | 7.65 | 360 | 234 | 132 | 0.08 | 30 | 35 | 135 | 0.03 |
| | Brembal | 7.65 | 410 | 266.5 | 135 | 0.3 | 43 | 38 | 141 | 0.02 |

ppm: Part per million = mg/L, EC: Electrical conductivity, $\mu\text{m/cm}$: $\mu\text{mohs / cm}$, TDS: Total dissolved salts, TH: Total hardness, Cl₂: Chlorine, Cl⁻: Chloride, SO₄²⁻: Sulfate, Alkal: alkalinity

the permissible limits of WHO⁶ for samples of up-streams of shark, Mit-khamis, sherbin, Bosat, Bilqas, El-Gamalia, El-Manzala stations and Abo-dawoud compact unit. Water Al concentrations were within the permissible limits of EMH⁵, however they equal to and/or exceeded the permissible limits of WHO⁶ for samples of Talkha, Sherbin, Bosat stations and network of Bosat station, respectively. Other measured parameters were within the permissible limits. Table 3 shows the values of detected heavy metals, Fe, Mn, Pb, Ni, Cr, Zn, Cu, Co and Cd in surface water samples. The concentration of Cd exceeded the permissible limits of EMH⁵ and WHO⁶ in the samples of up-streams of shark and Mit-Khamis stations, net-

works of Mit-Antar, Demera, Bosat, Bilqas, El-satamony, El-Gamalia, Mit-asim and Bilqas station. The concentration of Ni was more than the permissible limits of EMH⁵ and WHO⁶ in the network samples of Shoha, Bosat and El-Gamalia as well as up-stream of Bosat station. Lead concentrations in the network water samples of Shoha, Mit-Antar, Demera and Nabaru exceeded the permissible limits of EMH⁵ and WHO⁶.

Discussion

An estimate of Witt, based on WHO reports suggests that 80% of all human illness in the developing countries is associated with polluted water.⁷

Table 3: Results of some heavy metals in ppm of (up-stream, main station and network) samples

| No | District/sample name | Cd | Cu | Zn | Cr | Ni | Pb | Mn | Fe | Co |
|-----|--|--------------|------------|------------|-------------|-------------|-------------|------------|------------|-------------|
| | Permissible limit of WHO (2008) | 0.003 | 2.0 | 3.0 | 0.05 | 0.07 | 0.01 | 0.4 | 0.3 | 0.05 |
| | Permissible limit of EMH (2007) | 0.003 | 2.0 | 3.0 | 0.05 | 0.02 | 0.01 | 0.4 | 0.3 | 0.05 |
| | El-Mansoura district | | | | | | | | | |
| | Up-stream of shark station | 0.004 | 0.0 | 0.0 | 0.004 | 0.011 | 0.001 | 0.0 | 0.0 | 0.002 |
| | Main shark station | 0.002 | 0.0 | 0.0 | 0.002 | 0.008 | 0.001 | 0.0 | 0.0 | 0.002 |
| | Up-stream of Mit-khamis station | 0.006 | 0.0 | 0.1 | 0.004 | 0.006 | 0.003 | 0.0 | 0.01 | 0.001 |
| I | Network of Mit-khamis station | 0.003 | 0.0 | 0.0 | 0.001 | 0.003 | 0.0 | 0.0 | 0.0 | 0.001 |
| | Network of Mahalet damana | 0.001 | 0.0 | 0.0 | 0.005 | 0.006 | 0.002 | 0.0 | 0.0 | 0.001 |
| | Shoha Station | 0.0 | 0.0 | 0.1 | 0.008 | 0.009 | 0.008 | 0.01 | 0.01 | 0.0 |
| | Network of Shoha | 0.0 | 0.0 | 0.1 | 0.008 | 0.022 | 0.016 | 0.01 | 0.02 | 0.0 |
| | Network of El-Rydanian | 0.001 | 0.0 | 0.0 | 0.005 | 0.002 | 0.0 | 0.0 | 0.0 | 0.001 |
| | Talkha district | | | | | | | | | |
| II | Network of Mit-antar | 0.011 | 0.0 | 0.0 | 0.008 | 0.013 | 0.033 | 0.0 | 0.0 | 0.0 |
| | Network of Demera | 0.032 | 0.0 | 0.0 | 0.003 | 0.007 | 0.025 | 0.0 | 0.0 | 0.004 |
| | Dekernis district | | | | | | | | | |
| III | Network of Mit-Faris station | 0.0 | 0.0 | 0.0 | 0.001 | 0.011 | 0.003 | 0.0 | 0.0 | 0.001 |
| | Sherbin district | | | | | | | | | |
| IV | Network of Sherbin station | 0.001 | 0.0 | 0.0 | 0.006 | 0.008 | 0.002 | 0.0 | 0.0 | 0.001 |
| | Up-stream of Bosat station | 0.001 | 0.0 | 0.0 | 0.003 | 0.021 | 0.0 | 0.0 | 0.0 | 0.001 |
| | Network of Bosat station | 0.005 | 0.0 | 0.0 | 0.003 | 0.022 | 0.0 | 0.0 | 0.0 | 0.001 |
| | Bilqas district | | | | | | | | | |
| V | Main station of Bilqas | 0.005 | 0.0 | 0.0 | 0.001 | 0.009 | 0.0 | 0.0 | 0.0 | 0.001 |
| | Network of Bilqas station | 0.007 | 0.0 | 0.0 | 0.005 | 0.005 | 0.0 | 0.0 | 0.0 | 0.001 |
| | Network of El-satamony | 0.007 | 0.0 | 0.0 | 0.0 | 0.010 | 0.003 | 0.0 | 0.0 | 0.002 |

Continued

Table 3: Results of some heavy metals in ppm of (up-stream, main station and network) samples

| No | District/sample name | Cd | Cu | Zn | Cr | Ni | Pb | Mn | Fe | Co |
|------|--|--------------|------------|------------|-------------|-------------|-------------|------------|------------|-------------|
| | Permissible limit of WHO (2008) | 0.003 | 2.0 | 3.0 | 0.05 | 0.07 | 0.01 | 0.4 | 0.3 | 0.05 |
| | Permissible limit of EMH (2007) | 0.003 | 2.0 | 3.0 | 0.05 | 0.07 | 0.01 | 0.4 | 0.3 | 0.05 |
| VI | El-Gamalia district | | | | | | | | | |
| | Network of El-Gamalia station | 0.010 | 0.0 | 0.0 | 0.001 | 0.021 | 0.002 | 0.0 | 0.0 | 0.001 |
| VII | El-Sinbillawin district | | | | | | | | | |
| | Network of El-Sinbillawin station | 0.002 | 0.0 | 0.0 | 0.0 | 0.008 | 0.004 | 0.0 | 0.0 | 0.002 |
| VIII | El-Manzala district | | | | | | | | | |
| | Network of El-Manzala station | 0.003 | 0.0 | 0.0 | 0.001 | 0.009 | 0.003 | 0.0 | 0.02 | 0.002 |
| | Network of El-Mawaged | 0.002 | 0.0 | 0.01 | 0.003 | 0.008 | 0.0 | 0.0 | 0.01 | 0.001 |
| IX | Nabaru district | | | | | | | | | |
| | Network of Nabaru | 0.001 | 0.0 | 0.0 | 0.004 | 0.006 | 0.038 | 0.0 | 0.0 | 0.0 |
| X | Temy el-amdid district | | | | | | | | | |
| | Network of Abo-dawoud | 0.003 | 0.0 | 0.0 | 0.003 | 0.008 | 0.001 | 0.0 | 0.01 | 0.0 |
| XI | Minyet el-nasr district | | | | | | | | | |
| | Network of Mit-asim | 0.005 | 0.0 | 0.0 | 0.0 | 0.018 | 0.003 | 0.0 | 0.01 | 0.001 |
| | Network of Brembal | 0.001 | 0.0 | 0.002 | 0.0 | 0.005 | 0.0 | 0.0 | 0.0 | 0.001 |
| XII | Aga district | | | | | | | | | |
| | Network of Nawasa | 0.001 | 0.0 | 0.0 | 0.001 | 0.001 | 0.004 | 0.0 | 0.0 | 0.015 |

The pH values play an important role in the interaction between heavy metals and many parameters. Toxicity with heavy metals can increase with basic pH values. The abnormality observed in water Al concentrations in the region is probably attributed to addition of excessive Al-alum in these locations. There are many health problems reported in association with aluminum in water. For example, senile dementia is associated with high concentration of Al in water.⁸ Water with high concentration of Al cannot be used for hemodialysis too.⁸ Concerning on the heavy metals which have overload concentra-

tions in the water samples and their environmental impact on human health.

Primarily, Cd is found in surface water as a pollutant from industries such as electroplating. Water concentration of Cd >0.01 mg/L, like in network samples of Mit-Antar and Demera, has the potential to cause anemia, growth retardation and hypertension.⁹ Webb reported that the geochemical effects of Cd on human health appear as bone and renal failure in populations drinking a contaminated water.¹⁰

Ni though an essential trace element is toxic in large amounts. Toxicity with Ni is

TAKE-HOME MESSAGE

- There are many health problems reported in association with aluminum in water.
- Cadmium is found in surface water as a pollutant from industries such as electroplating.
- Nickel though an essential trace element, is toxic in large amounts.
- Lead is a non-essential and highly toxic cumulative trace element in man.
- Continuous chemical analysis of surface drinking water is required to ensure that it is suitable for domestic purposes.
- Aluminum, cadmium, nickel and lead concentrations in surface drinking water samples of Dakahlia Governorate are higher than the permissible limits of EMH (2007) and WHO (2008).

enhanced in presence of Co, Cu, Fe, and Zn in drinking water. Kaaber, *et al.*, reported worsening of eczema for those exposed to high levels of Ni.^{11,12}

Pb is a non-essential and highly toxic cumulative trace element in man. It is present in tap water to some extent as a result of its dissolution from natural sources; Pb is primarily from household plumbing systems containing lead in pipes, solder, fittings or the service connections to homes. Levels of Pb in drinking water sampled at the source are usually <5 µg/L. However, tap water taken from homes where lead is present in the plumbing can contain levels up to 100 µg/L.¹³ The presence of lead in water usually indicates contamination from metallurgical wastes or from lead-containing industries. Lead concentrations in the network water

samples of Shoha, Mit-Antar, Demera and Nabaru exceeded the permissible limits of EMH⁵ and WHO⁶. Lead after absorption, enters the blood where it is rapidly taken-up by the erythrocytes. High concentration of lead in the body can cause death or permanent damage to the central nervous system, the brain and kidney.¹⁴ The damage commonly results in behavior and learning problems. Lead crosses the placenta easily throughout gestation¹⁵ and is usually associated with poor pregnancy and neonatal outcomes. Other heavy metals are within the permissible limits.

Considering the current situation, continuous chemical analysis of surface drinking water is required to ensure that it is suitable for domestic purposes. Since most of the measured heavy metals are most likely originate from steel, plastics and batteries industries working in the region, we believe that activities of these industries must be stopped or at least limited in urban zones. We should also emphasize on using appropriate materials and engineering practices to minimize plumbing solvency in water treatment and water distribution systems.

Conflicts of Interest: None declared.

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