

Levels and ecological risk of pharmaceuticals in River Sosiani, Kenya

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Abstract The continued frequent detection of pharmaceuticals in the environment is of major concern due to potential human and ecological risks. This study evaluated 30 antibiotics from 8 classes: sulphonamides (SAs), penicillins (PNs), fluoroquinolones (FQs), macrolides (MLs), lincosamides (LINs), nitroimidazoles (NIs), diaminopyrimidines (DAPs), salfones and 4 anthelmintics benzimidazoles (BZs) in surface water and sediments from River Sosiani in Eldoret, Kenya. Samples were collected during the wet and dry seasons and subjected to solid phase extraction using HLB cartridges. A liquid chromatography tandem mass spectrometry (LC–MS/MS) method was used for the simultaneous quantification

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Faculty of Science and Technology, Department of Chemistry, Multimedia University of Kenya, P.O. Box, 15653-00503 Nairobi, Kenya e-mail: ngugianastasiah@gmail.com; angigi@mmu.ac.ke of the compounds. Chromatographic separation was on a reversed-phase Zorkax Eclipse Plus C18 column eluted in a gradient program and compounds detected by mass spectrometer operated in a positive electrospray ionization (+ESI) mode. Twenty-eight antibiotics were detected in water where 22 had a 100% detection frequency and the remaining 4 showed detection frequencies ranging from 5 to 47%. Three BZs had a 100% detection frequency. Detectable concentrations of the pharmaceuticals in water ranged between 0.1 and 247 ng L^{-1} and 0.01 and 974 µg kg⁻¹ in the sediments. The sulfonamide, sulfamethoxazole, had the highest concentration in water (247 ng L^{-1}), whereas penicillin G showed the highest concentrations in sediments (414-974 µg kg⁻¹). Quantified pharmaceuticals decreased in the order SAs>DAPs >FQs>ATs>PNs \approx MCs \approx LNs>NIs in water, and followed the order PNs > BZs > FQs > MLs > DAPs \approx LNs > NIs > SAs in sediments. Risk quotients (RQ_w) showed that sulfamethoxazole and ciprofloxacin were of high ecological risk in the surface water (RQ_w values of 1.11 and 3.24, respectively), whereas penicillin V, ampicillin, penicillin G, norfloxacin, enrofloxacin, erythromycin, tylosin, and lincomycin were of medium ecological risk in the aquatic system. The findings show high prevalence of pharmaceuticals in surface water and sediments and are therefore potential ecological hazards. Such information is vital when devising mitigation strategies.

Keywords Antibiotics \cdot Anthelmintics \cdot Surface water \cdot Sediments \cdot LC-MS/MS \cdot Ecological risks

Introduction

Antibiotics are natural, synthetic, and semi-synthetic compounds which show antimicrobial activities on bacteria or other single-celled microorganisms (Catteau et al., 2018). Antibiotics have revolutionized medicine in many respects, and countless lives have been saved; their discovery was a turning point in human history (Davies & Davies, 2010). They represent one of the largest therapeutic categories used in the treatment of infectious diseases caused by bacteria (Schmieder & Edwards, 2012). Antibiotic drugs are predominantly used to treat bacterial diseases in human therapy and as veterinary medicines to prevent and treat bacterial diseases in animal husbandry (Kümmerer, 2009). Besides these fundamental applications of antibiotics in treatment of certain types of diseases in animals and humans, a significant portion is used in animal feed as supplements to promote animal growth (Chattopadhyay, 2014; Economou & Gousia, 2015; Poole & Sheffield, 2013), more so in developing countries where it is harder to successfully regulate the use of antibiotics (Eagar et al., 2012; Van et al., 2020). This broadness of applications has caused an increase in the consumption of antibiotics (Kraemer et al., 2019; Pokharel et al., 2020).

The residual antibiotics from human and animal use can enter the environment via various pathways, including wastewater effluent discharge, runoff from land to which agricultural or human waste has been applied, and leaching (Bottoni et al., 2010; Faleye et al., 2018; Polianciuc et al., 2020; Sodhi et al., 2021). Antibiotics might also be added to the environment through improper disposal of unused and expired antibiotics, where they end up in wastewater, landfills, water supplies, and drains (Anwar et al., 2020; Kotwani et al., 2021). Domestic effluents are considered a major contributor of antibiotic contamination of surface and wastewaters (Zheng et al., 2012), but effluents from pharmaceutical industries (Kotwani et al., 2021) and hospitals' effluents are also of great concern (Bansal, 2019; Brown et al., 2006; Ngigi et al., 2020). Furthermore, the intensive practice of aquaculture for food production can lead to inadvertent introduction of antibiotics into surrounding surface waters (Hoa et al., 2011; Zheng et al., 2012).

Antibiotics are often partially metabolized after administration, and a significant portion of the antibiotic can be excreted as the parent compound or in conjugated forms that can be converted back to the parent antibiotic. For example, fluoroquinolones, whose therapeutic prescription in human medicine is between 300 and 600 mg per day, are almost all excreted as unchanged compounds in urine and are, consequently, discharged into hospital sewage or municipal wastewater (Lindberg et al., 2004). This ultimately contributes to residual antibiotics in recipient waters. Thus, sewage sludge and manure, used as fertilizer in agricultural land, are often contaminated with antibiotics (Thiele-Bruhn, 2003). Subsequent runoffs from such contaminated agricultural land transfer some of the residues into environmental water systems such as surface water.

Therefore, the continued intensive use of antibiotics in livestock production and human medication has led to their frequent detection in different environmental matrices. Antibiotics have been reported in hospital wastewater effluents (Anwar et al., 2020; Aydin et al., 2019; Bansal, 2019; Diwan et al., 2009; Lindberg et al., 2004; Ngigi et al., 2020; Yao et al., 2021), wastewater treatment plant (WWTP) effluents (Golchin et al., 2021; Kortesmäki et al., 2020; Ngigi et al., 2020; Rodriguez-Mozaz et al., 2020), WWTP biosolids, soil, surface waters, groundwater (Burke et al., 2016; Clarke & Smith, 2011; Hu et al., 2018; Li et al., 2021; Martín et al., 2015; Yang & Carlson, 2003), sediments, biota and drinking water (Hanna et al., 2018; Li et al., 2014; Meng et al., 2021; Wang et al., 2016), and in different forms of natural waters systems (Litynska et al., 2021) at ng L^{-1} to low $\mu g L^{-1}$ levels. It is therefore important to continuously monitor the residual antibiotics in WWTPs and in surface water. There are few studies on surveillance of antibiotic residues in the environment in Kenya. For river surface water, antibiotics from different classes were reported at varying concentrations including the sulphonamides sulfamethoxazole, sulfamethazine, and sulfadiazine at concentrations of up to 142.6, 0.47, and 0.84 μ g L⁻¹, respectively; β -lactam (penicillins) including amoxicillin (up to 9 μ g L⁻¹), ampicillin (up to 0.9 μ g L⁻¹), and penicillin G, dicloxacillin, and nafcillin (<1.0 μ g L⁻¹); the fluoroquinolones ciprofloxacin and norfloxacin (up to 4.9 and 2.8 μ g L⁻¹); and trimethoprim (up to 9.5 μ g L⁻¹) (K'oreje et al., 2012; Kairigo et al., 2020a, b; Kimosop et al., 2016; Ngigi et al., 2020; Opanga, 2018). Other environmental matrices studied include river sediments (Kairigo et al., 2020b; Kandie et al., 2020; Kimosop et al., 2016), hospital wastewater, and lagoons and WWTPs (Kimosop et al., 2016; Ngigi et al., 2020).

Establishing residual antibiotics in water sediments, surface and hospital wastewater is important for baseline information, generation of knowledge, bridging existing knowledge gaps, and in policy formulations. This study investigated the occurrence of 4 anthelmintics benzimidazoles (BZs) and 28 antibiotics from the classes: sulphonamides (SAs), β-lactams-penicillins (PNs), Fluoroquinolones (FQs), macrolides (MLs), lincosamides (LINs), nitroimidazoles (NIs), and two other antibiotics dapsone (a sulfone) and trimethoprim (a Diaminopyrimidine) in sediments and surface water from River Sosiani in Eldoret Municipality. Kenya, as one of the developing countries, has few reports on the status of pharmaceuticals contamination of surface and wastewaters. Environmental risks can only be assessed when the background contamination levels have been established.

Materials and methods

Chemicals and reagents

Standard pharmaceutical reference compounds (>98% purity) and high-performance liquid chromatographygrade water (LC water) were obtained from Merck through Scientific Laboratory Supplies Ltd. (Kenya

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Branch). Table 1 gives the list of the compounds and abbreviations, while table S1 (supplementary material) gives the physicochemical properties of the compounds used in this study. High-performance liquid chromatography-grade methanol (MeOH) and acetonitrile (ACN); formic acid (HCOOH) and ammonium formate (NH₄HCO₂) were purchased from Kobian Scientific Ltd. (Nairobi, Kenya). Oasis hydrophilic–lipophilic balance cartridges (HLB, 200 mg/6 mL; 60 mg, 3 mL) and glass microfiber filters (0.7, 0.45, 0.22 μ m) were obtained from Milford, MA (Waters, USA). All other reagents were of analytical grade and were purchased from Kobian Scientific Ltd. Individual standard stock solutions were prepared at a concentration of 1 g/L, in either methanol or acetonitrile and stored at 4 °C.

Study area and sampling

The study was conducted in Uasin Gishu County, within Eldoret municipality (Fig. 1) in Kenya. Surface water samples and water sediments were collected from River Sosiani (which traverses through the town) in Eldoret Municipality during the wet season (June, 2020) and the dry season (February, 2021). A judgemental random sampling method was used whereby 10 sampling points were identified (Table 2) in the suburban and the urban sections of the river

Table 1 List of compounds used in the study

Class of antibiotics				
Sulfonamides (SAs)	Sulfaguanidine (SFG)	Sulfamonomethoxine (SMM)	Sulfachlorpyridazine (SCP)	Sulfadimethoxine (SDMX)
	Sulfadiazine (SDZ)	Sulfamethazine (SMZ)	Sulfamethoxazole (SMXZ)	Sulfadoxine (SDX)
	Sulfathiazole (STZ)	Sulfamethoxypyridazine (SMPZ)	Sulfisoxazole (SXZ)	
	Sulfamethizole (SMTZ)	Sulfapyridine (SPD)	Sulfaquinoxaline (SFQ)	
β-lactams (penicillins	Penicillin V (PEV)	Amoxicillin (AMX)	Cloxacillin (CLX)	Nafcillin (NAF)
(PNs)	Ampicillin (AMP)	Penicillin G (PEG)	Dicloxacillin (DCX)	
Fluoroquinolones (FQs)	Norfloxacin (NOR)	Ciprofloxacin (CIP)	Enrofloxacin (ENR)	
Macrolides (MLs)	Erythromycin (ERY)	Tylosin (TYL)		
Lincosamides (LNs)	Lincomycin (LIN)			
Nitroimidazoles (NIs)	Metronidazole (MET)			
Diaminopyrimidines (DAPs)	Trimethoprim (TMP)			
Salfone	Dapsone (DPS)			
Anthelmintics	1 · · ·			
Benzimidazoles (BZs)	Oxfendazole (OXF)	Mebendazole (MEB)	Flubendazole (FLB)	Albendazole (ALB)



Fig. 1 Location of sampling points in the study area

in close proximity to urban settlements, hospitals, and the city's sewer system or where wastewater discharges occur followed by random sampling from each selected site. Water samples were collected in 2.5 L amber glass bottles that had been pre-cleaned successively with saturated MeOH-EDTA solution, rinsed with distilled water, and then dried in an oven at 100 °C. Prior to sample collection, the bottles were pre-rinsed three times with river water. Five 0.5-L grab water samples were collected randomly at each sampling point (away from the edges of the river) and combined to form a composite 2.5 L sample (in duplicate). The temperature and pH were recorded in situ. The pH of the water ranged from 7.0 to 8.0, with

 Table 2
 Sampling locations and some parameters of water samples

Sampling point	Sample code	Coordinates	рН	Altitude (above sea level) (ft)	Temperature (°C)	Location/major anthropogenic activity
1	SP1	0° 30' 33.5" N35° 16' 42.8" E	7.4	200	19.3	Forest
2	SP2	0° 30' 32.7" N35° 16' 43.4" E	7.2	50	22.6	Hospital
3	SP3	0° 30' 34.8" N35° 16' 43.1" E	7.2	50	21.8	Hospital
4	SP4	0° 30' 42.0" N35° 16' 33.5" E	7.8	100	22.7	Hospital
5	SP5	0° 30' 50.7" N35° 16' 47.1 " E	7.8	200	23.7	Bridge
6	SP6	0° 30' 45.7" N35° 16' 34.6" E	7.4	200	22.8	Hospital
7	SP7	0° 30' 42.9" N35° 16' 39.3" E	7.8	83.3	21.5	Hospital
8	SP8	0° 30' 41.0" N35° 16' 39.0" E	7.5	200	21.5	Forest
9	SP9	0° 30′ 31″ N 35° 17′ 2″ E	7.9	83.3	20.4	Residential
10	SP10	0° 30' 31" N 35° 16' 59" E	7.8	83.3	19.7	Bridge

majority of the samples being neutral. Sodium azide (0.5 g L^{-1}) was added into each sample bottle immediately after sampling to inhibit potential biodegradation. Samples were placed in cooler box during transportation, acidified with HCl to a pH of 3 and stored at 4 °C in a cold room prior to extraction. Ten grab sediment samples (approximately 200 g) were also collected (in duplicate), coded as SP1 to SP10, and transported together with the water samples, where they were stored at -20 °C in a cold room prior to extraction.

Sample preparation

One-liter subsamples (from the composite samples) were filtered sequentially using 0.7 µm and 0.45 µm Millipore filter prior to SPE clean-up. The Oasis hydrophilic-lipophilic balance (HLB) cartridges (200 mg, 6 mL) were preconditioned with 6 mL MeOH followed by 6 mL LC-water. Each 1-L water sample (in triplicate) was then passed through the cartridge at a flow rate of 8 mL/min using an SPE vacuum Manifold (MilliporeSigmaTM SupelcoTM Visiprep, Thermo Fisher Scientific), washed with 10% MeOH in LC-water and air-dried for 10 min. The cartridges were eluted with 3 mL of 70:30 v/v MeOH-ACN solution and reduced to a volume of 100 µL under a gentle flow of nitrogen. The sample was reconstituted to 1 mL with a 2:1 (v/v)LC-Water-ACN solution containing 5 mM NH₄HCO₂ (acidified with 0.1% HCOOH), filtered through a 0.22µm glass membrane filter and stored at 4 °C prior to LC-MS/MS measurement.

For the sediment samples, a 2-g aliquot of sample was weighed into a 50-mL centrifuge tube. Six milliliter of MeOH: ACN: H₂O: water (60: 20: 20) solvent mixture was added to the tube followed by vortexing for one minute. The mixture was then sonicated for 20 min and centrifuged at 4500 rpm for 10 min, after which the supernatant was collected in a 15-mL glass test tube. A repeat extraction using 6 mL of 100% methanol was done and extracts were pooled in the 15-mL tube. The pooled extract was concentrated using a rotary evaporator to approximately 1 mL and reconstituted to 10 mL using LC-water. Extracts were passed through 0.45-µm glass microfilters and cleaned-up by SPE method using Oasis (HLB) cartridges (60 mg, 3 mL). The SPE procedure was same as that of the water samples, using 3 mL of preconditioning, washing, and eluting solutions.

LC-MS/MS analysis

Samples were analyzed using Agilent LC-1290 infinity II system (Germany) coupled to an API 6460c triple Quad Mass Spectrometer (Applied Biosystems/ MDS Sciex Instruments, Toronto, Canada) system equipped with a Zorbax Eclipse Plus C18 RRHD, 50 mm × 2.1 mm, 1.8 µm column and a guard column $(3.0 \times 4 \text{ mm})$ of the same material. The binary solvent gradient elution program (Solvent A: 0.1% aqueous HCOOH, 5 mM ammonium formate and Solvent B: 0.1% HCOOH acetonitrile) of 23 min was set up as follows: 0 min, 0% B; 1 min, 5% B; 3 min, 10% B; 7 min, 20% B; 11 min, 30% B; 15 min, 40% B; 17 min, 50% B; 19 min, 70% B; 21 min, 80% B; and 23 min, 5% B. The oven temperature was set at 35 °C, a flow rate of 0.30 mL min⁻¹, and an injection volume of 10 µL. A post-time of 3 min was set between sample runs for the column re-equilibration. Mass spectrometer (MS) was operated in positive electrospray ionization (+ESI) mode with the following source parameters: gas temperature 325 °C, gas flow 1 L min⁻¹, and nebulizer gas 45 psi capillary voltage of 4000 V. Precursor ion and two or three product ions were used for compound identification (Supplementary Table S2).

Method evaluation and validation

The SPE clean-up method was evaluated by spiking mixed standards to 1-L blank water samples at two concentrations (0.01 and 0.5 μ g L⁻¹) and subjecting the samples to the same SPE procedure as the samples. One-gram triplicate sediment samples were also spiked to a final concentration of 0.1 and 0.5 μ g kg⁻¹ and subjected to similar extraction and SPE clean-up procedure as the sediment samples. Recoveries (%) were determined for each of the analyte compound as the ratio of quantified amount to that of the spiked amount multiplied by 100%. Repeat blank sample spike experiments (in water) were conducted in different days to evaluate the intra-day and inter-day method precision and accuracy of recoveries for representative test pharmaceuticals from different classes (Table S3, supplementary material).

Reference standard compounds of concentrations 0.1–2 $\mu g~mL^{-1}$ in 2:1 (v/v) LC-Water-ACN solution containing 5 mM NH_4HCO_2 (acidified with 0.1% HCOOH) were prepared and injected into the

LC–MS/MS to determine linearity of the quantification method and as external calibration standards. A set of six replicates of lowest calibration mixed standard were injected into LC–MS/MS instrument and used to determine the limit of detection (LOD) and limit of quantification (LOQ). LOD and LOQ were evaluated as; LOD= $(3.3 \times \sigma/s)$, while LOQ= $(10 \times \sigma/s)$, where σ is standard deviation of the replicates and *s* is the slope. There was good precision among the replicates for each compound, and this reproducibility confirmed ruggedness of method.

The determined method performance parameters including the linearity, accuracy, and repeatability expressed as the correlation coefficient (R^2) of calibration curves,% recoveries, LOD, LOQ, and standard deviation for replicate measurements for water samples, and spiked samples are presented in Tables 3 and S3 (supplementary material). Linearity was good ($R^2 > 0.93$) for all test compounds. The% recoveries ranged from 83 to 114% for water samples and 80–108% for the sediments. The LOD of the adopted method ranged from 0.003 to 0.317 ng L⁻¹ whereas the LOQ range was 0.010–0.908 ng L⁻¹. All values reported were above the LOQ for each corresponding analyte compound. The method was therefore robust enough for the study.

Ecological risk assessment

The potential ecological risk of the pharmaceutical residues in water was evaluated using risk quotient RQ_w, which was evaluated based on maximum measured environmental concentration (MEC) to the predicted no-effect concentration (PNEC, ng L^{-1}) using Eq. (1) (Yin, 2021).

$$RQ_{w} = \frac{MEC}{PNEC_{w}}$$
(1)

 $PNEC_w$ (maximum drug concentration with no adverse effect on the microorganisms or the ecosystem in the environment) was determined using the evaluation (or assessment) factor, AF (Eq. 2).

$$PNEC_{w} = \frac{NOEC \text{ or } EC_{50}}{AF}$$
(2)

where EC_{50} is the half-maximum effect concentration (ng L⁻¹). EC_{50} and $NOEC_{50}$ values (for the most sensitive species) were obtained from literature (Chen et al., 2021a; Oh et al., 2004; Sharma et al., 2021). NOEC₅₀ values were used in cases where EC₅₀ could not be obtained (Table S7, supplementary material). An AF value of 1,000 was used when acute toxicity (EC₅₀) data was used, and 100 when chronic toxicity data, NOEC₅₀ was used (Jiang et al., 2014). From the commonly used criteria of ranking the RQ values, RQ range of 0.01 to 0.1 is considered as low risk, 0.1 > RQ < 1 medium risk, and RQ > 1 as high risk, and this criterion was adopted in this study (Yan et al., 2013).

Data analysis

Descriptive and other data analysis were done using Microsoft Excel 2016 (Apple Inc., USA) computer software. Pearson's correlation (r, $\rho = 0.05$) was used for the comparison of pharmaceutical residues in river surface water and in sediments. Statistical comparison of the intra- and inter-day recoveries was done to evaluate the precision and accuracy (% coefficient of variation, CV, and% accuracy) of the method. Pseudo-partitioning coefficients $(K_{n,s})$ of test compounds between water and sediments were used to estimate the sorption capacity of these compounds and were calculated using the equilibrium: $K_{p,s} = C_s/$ $C_{\rm w}$, where $C_{\rm w}$ is the mean concentration of the target compound in the water sample ($\mu g L^{-1}$), and C_s is the mean concentration of the target compound in the sediment sample (ng g^{-1}) (Chen et al., 2021b).

Results and discussion

Residual pharmaceuticals in water

Water samples were randomly obtained from 10 selected sites in Sosiani river (Eldoret) and analyzed for antibiotic residues. From a total of 34 compounds that were investigated, varied amounts of residues were detected for 28 compounds is surface water, representing 79% of analytes. The six compounds that were not detectable included sulfaguanidine, sulfamonomethoxine, sulfachlorpyridazine, sulfaquinoxaline, dapsone, and flubendazole. Detectable amounts of SAs ranged from 0.08 to 247.0, PNs from 0.19 to 10.37, QNs from 13.68 to 56.02, MCs 3.32 to 9.08, LNs (LIC) from 3.67 to 7.77, NIs (MET) from 3.11 to 5.56 and TMP ranged from 0.17 to 67.32 ng L⁻¹. The

Compound	Calibration equation	R^2	% recovery in water	% recovery in sediments	$LOD \ (ng \ L^{-1})$	LOQ (ng L ⁻¹)
SFG	y = 1255.24x + 1813.70	0.999	90.00 ± 8.01	93.44 ± 9.48	0.028	0.084
SDZ	y = 2084.72x + 386.77	0.998	113.8 ± 3.05	108.00 ± 12.73	0.023	0.069
STZ	y = 1578.08x + 6322.26	0.988	98.44 ± 8.00	98.88 ± 6.90	0.119	0.359
SMTZ	y = 539.11x + 345.51	0.996	92.91 ± 7.88	89.47 ± 8.68	0.006	0.019
SMM	y = 5433.86x + 19,791.77	0.995	98.70 ± 3.25	99.77 ± 3.45	0.006	0.018
SMZ	y = 2348.55x + 9297.53	0.991	104.84 ± 11.08	100.23 ± 7.87	0.014	0.043
SMPZ	y = 5502.77x + 1405.27	0.997	93.98±4.69	80.07 ± 4.81	0.006	0.018
SPD	y = 616.62x + 1494.97	0.999	98.20 ± 2.07	93.33 ± 4.71	0.054	0.162
SCP	y = 2014.77x + 7861.64	0.990	87.57 ± 5.52	99.44 ± 2.51	0.016	0.050
SMXZ	y = 1361.40x + 5067.49	0.989	109.03 ± 8.48	100.45 ± 4.41	0.056	0.170
SXZ	y = 1104.31x + 2689.19	0.998	98.70 ± 3.25	97.88 ± 2.66	0.040	0.122
SFQ	y = 1476.14x + 12,278.26	0.982	97.70 ± 6.18	97.45 ± 0.31	0.022	0.068
SDMX	y = 6876.99x + 10,819.36	0.999	101.87 ± 13.95	81.09 ± 1.54	0.005	0.016
SDX	y = 3487.36x + 10,381.48	0.999	100.69 ± 4.28	97.44 ± 0.32	0.014	0.041
DPS	y = 553.29x + 1058.94	0.999	97.30 ± 7.50	93.43 ± 5.79	0.006	0.018
NOR	y = 857.10x - 18,865.35	0.928	96.53 ± 5.47	94.63 ± 5.60	0.092	0.280
CIP	y = 1587.47x - 21,575.60	0.984	101.33 ± 6.60	97.69 ± 0.44	0.044	0.133
ENR	y = 1323.60x - 24,880.26	0.955	95.07 ± 7.64	97.78 ± 2.52	0.109	0.330
PEV	y = 135.53x - 385.45	0.998	96.77±5.33	98.16 ± 5.88	0.317	0.960
AMP	y = 168.78x + 159.37	0.993	88.20 ± 4.05	82.84 ± 2.60	0.020	0.059
AMX	y = 392.24x - 460.66	0.998	80.79 ± 1.12	82.80 ± 1.61	0.014	0.043
PEG	y = 49.86x + 140.25	0.997	91.68 ± 10.86	98.76 ± 4.37	0.113	0.341
CLX	y = 14.89x - 29.00	0.997	85.73 ± 8.11	90.65 ± 7.51	0.300	0.908
DCX	y = 2091.96x - 6421.16	0.999	104.01 ± 12.74	95.87 ± 4.53	0.089	0.270
NAF	y = 7026.26x - 24,981.86	0.999	103.06 ± 6.68	90.14 ± 9.63	0.008	0.023
ERY	y = 2862.05x - 11,347.62	0.993	83.01 ± 4.26	100.37 ± 10.89	0.082	0.247
TY L	y = 1883.39x - 15,649.47	0.993	98.98 ± 7.51	96.63 ± 4.20	0.018	0.055
LIC	y = 3580.88x - 20,237.15	0.998	91.71 ± 6.19	96.22 ± 5.78	0.015	0.045
MET	y = 1963.60x + 6079.99	0.999	88.98 ± 5.62	97.37 ± 4.25	0.053	0.162
TMP	y = 1627.21x - 31.47	0.998	86.47 ± 4.43	81.04 ± 7.13	0.027	0.082
OXF	y = 4624.85x - 8946.04	0.999	98.27 ± 2.92	98.95 ± 2.76	0.003	0.010
MEB	y = 14,067.99x - 37,179.38	0.999	89.07 ± 6.23	98.84 ± 4.01	0.011	0.035
FLB	y = 8944.22x + 2596.94	0.999	92.04 ± 3.83	94.27 ± 7.45	0.004	0.011
ALB	y = 9018.73x - 24,555.95	0.999	92.44 ± 10.99	93.20 ± 10.65	0.014	0.043

amount of determined BZs (OFX, MEB, and ALB) ranged from 1.97 to 21.39 ng L⁻¹. Therefore, the concentrations (based on maximum amount) decreased in the order SAs>TMP>FQs>ATs>PNs \approx MCs \approx LNs>NIs (Tables 4, 5 and 6). Though the highest detected amount was that of a sulfonamide compound (SMXZ), the FQs were generally present in higher amounts than the other classes of compounds determined (Tables 4, S4a and b). The three FQs were detected at all sampling points, presenting a 100% frequency. AMP was detected at only one sampling point, SP4 (0.2 ng L⁻¹), a similar observation to that of STZ. PEG was detected at four sampling points (2% frequency), whereas the other two PNs (AMX and CLX) were detected at all sampling points. For all other detected compounds (other than SAs, FQs, and PNs), 100% detection frequency was observed. There was no observed trend in concentrations of detected compounds downstream. This suggests that dilution was not an influencing factor, and point source pollution may have played a major role, taking into account that the sampling area has several hospitals, residential areas (discharge of wastewater) and, to a small extent, some agricultural activities (Table 2). Sampling point SP4 recorded higher concentrations than other sites for most of the compounds, which strongly supports the idea of point source pollution as the site may be a recipient of large amounts of wastewater from a nearby hospital. Surface runoffs, from poorly managed domestic and hospital wastes and effluents, might have contributed to the pharmaceuticals in the investigated surface water.

Of the 10 SAs present in detectable amounts, SMXZ had the highest concentration of 247.0 ng L^{-1} followed by SMZ with 23.3 ng L^{-1} (Tables 4 and S4a-supplemantary material). The average concentration increased in the order SMPZ < SDZ < SMTZ <SDMT<SDX<SXZ<SPD<SMZ<SMXZ. This observed trend did not follow the order of solubility of the compounds in water, and was not significantly correlated (r=0.39, $\rho=0.05$) to the octanol-water coefficient (log K_{ow}). STZ was detected at only one sampling point (SP4) in the wet season at a concentration of 7.4 ng L^{-1} (Table 4). The frequency of detection of SAs varied among determined compounds and ranged from 0.05 to 100%. Considering the average amount for each compound at all sampling points, there was no discernable trend in the concentrations of SAs while comparing amounts in dry and wet seasons (supplementary Fig. S1a). This indicated contributions from point sources as well as surface runoffs. In general, SMTZ, SXZ, SDMT, and SDX recorded higher amounts in wet season (difference not significant), SPD and SXZ had higher amounts in dry season (difference significant for SPD, and significant for most sites for SXZ) whereas the concentrations of SMXZ and SMPZ were variable for the two seasons. It was noted that point sources contributed more to the concentrations of SDZ, SMTZ, SMZ, SMPZ, SPD, SMXZ, and SXZ during the dry season for points SP4 and SP6, and a similar observation for SDZ, SMZ, SMPZ, SPD, SMXZ, and SXZ for the points SP5, SP7, SP8, and SP9. Compared to other sampling points, SP4 had the highest concentrations of SDZ, SMTZ, SMZ, SMPZ, SMXZ, and SXZ in the dry season. Hospital and domestic effluents were the major sources of SAs in the sampled surface water (Table 2), going by the description of the study area.

Globally, reported concentrations of various SAs in surface water vary from non-detectable (nd) levels to 5320 ng L^{-1} for SMXZ (Chen & Zhou, 2014; Dinh et al., 2017; Kuang et al., 2020; Matongo et al., 2015; Ncube et al., 2021; Zhang et al., 2012), and up to 6192, 112.5, and 530.1 ng L^{-1} for SMZ, SDZ, and SPD, respectively (Chen & Zhou, 2014; Díaz-Cruz et al., 2008; Dinh et al., 2017; Yao et al., 2017). Also, a reported amount of up to 66.0 and 6.0 ng L^{-1} for SMM and SFQ, respectively, (Chen et al., 2018), and up to 47.5 and 15.6 ng L^{-1} STZ and SXZ, respectively (Li et al., 2021). However, high levels of SMXZ of 19.4 and 142.6 μ g L⁻¹ were reported for upstream and downstream of Mitheu river water in Kenya (Kairigo et al., 2020a). Other reported SA contaminations in surface water (Nairobi, Chania, Sagana, Kanyulu, and Mwania rivers) in Kenya are concentrations of up to 23.35, 0.47, and 0.84 μ g L⁻¹ of SMXZ, SMZ, and SDZ, respectively (K'oreje et al., 2012; Kairigo et al., 2020b; Ngigi et al., 2020), whereas values of $< 0.08 \ \mu g \ L^{-1}$ were determined for SDX and SCP (Ngigi et al., 2020).

Concentrations of the seven PNs ranged from 0.19 to 10.37 ng L^{-1} , with average amounts increasing in the AMP<AMX<PEG<CLX<PEV<DCX<N order AF (Tables 5 and S4b). A 100% frequency of detection was observed for five of the compounds except AMP (detected at sampling point SP4 during dry season only) and PEG (21% frequency). Also, notable observation was that the maximum concentrations of PEV, CLX, and DCX were obtained from sampling point SP4 during the wet season. Generally, concentrations of PEV, AMX, DCX, NAF, and CLX were high during the wet seasons (supplementary Fig. S1b), though the differences were not significant ($\rho = 0.05$) except AMX at SP9, DCX at SP4, and CLX at SP8 where the amounts were higher in dry the season). Among the PNs, AMP is among the most frequently detected antibiotic in aquatic systems, having a reported concentration of up to 13,800 ng L^{-1} in surface water (Dinh et al., 2017; Ncube et al., 2021). Others include AMX, PEG, and PEV at concentrations 200, 250, and 10 ng L^{-1} , respectively, in surface water (Dinh et al., 2017; Watkinson et al., 2009). In Kenya, the reported penicillin's in water include AMX at 900 ng L^{-1} in Mitheu river (Kairigo et al., 2020b), AMP, PEG, DCX, NAF, and oxacillin at a range of < 120-500 ng L⁻¹ (Ngigi et al., 2020) in Nairobi river.

Concentrations of the three FQs (100% frequency detection) ranged from 13.68 to 56.02 ng L^{-1} ,

Table 4 Conc	entrations of SA	s in surface wate	${ m tr} \ { m in} \ { m ng} \ { m L}^{-1}$							
Sample site	SDZ	STZ	SMTZ	SMZ	SMPZ	SPD	SMXZ	ZXZ	SDMT	SDX
SP1 W	nd	nd	0.94 (0.15)	nd	0.73 (0.13)	2.53 (0.06)	nd	2.51 (0.05)	1.89 (0.07)	3.34(0.11)
SP1 D	nd	nd	0.70~(0.06)	nd	0.33~(0.05)	2.92 (0.26)	pu	2.74 (0.07)	1.65 (0.07)	3.04(0.01)
SP2 W	nd	nd	0.99(0.13)	nd	0.33~(0.03)	2.61 (0.13)	pu	2.54 (0.09)	1.81 (0.02)	3.19(0.13)
SP2 D	nd	nd	0.70(0.00)	nd	0.33~(0.03)	2.80 (0.04)	pu	2.82 (0.35)	1.64 (0.02)	3.06(0.03)
SP3 W	nd	nd	0.98(0.26)	nd	0.44~(0.06)	2.52 (0.08)	pu	2.61 (0.11)	1.76 (0.02)	3.16(0.04)
SP3 D	nd	nd	0.78 (0.19)	nd	0.39~(0.05)	2.89 (0.20)	pu	2.63 (0.21)	1.66(0.00)	3.05(0.00)
SP4 W	0.44 (0.05)	7.38 (0.06)	0.83 (0.22)	0.73 (0.20)	0.71 (0.07)	2.82 (0.01)	10.11 (0.50)	2.55 (0.02)	3.69 (0.06)	3.20(0.03)
SP4 D	2.93 (0.28)	pu	1.13 (0.07)	16.33 (0.55)	1.16(0.09)	3.27 (0.11)	247.0 (5.59)	5.95 (0.55)	1.69(0.03)	3.17(0.07)
SP5 W	nd	nd	0.77 (0.14)	nd	0.45 (0.02)	2.61 (0.11)	9.00 (0.35)	2.65 (0.19)	1.74(0.10)	3.18(0.04)
SP5 D	nd	nd	0.70 (0.05)	0.80 (1.13)	0.48(0.09)	3.74 (0.15)	nd	4.72 (0.12)	1.68(0.08)	3.02(0.01)
SP6 W	nd	nd	0.75 (0.11)	nd	$0.36\ (0.03)$	2.45 (0.03)	pu	2.67 (0.23)	1.76 (0.03)	3.15(0.02)
SP6 D	0.24~(0.03)	nd	0.78 (0.15)	10.71 (0.82)	0.65(0.20)	5.83 (0.26)	4.21 (0.14)	2.70 (0.27)	1.63 (0.02)	3.10(0.09)
SP7 W	nd	nd	0.88 (0.06)	nd	$0.36\ (0.00)$	2.74 (0.32)	nd	2.69 (0.11)	1.71 (0.01)	3.10(0.04)
SP7 D	0.08 (0.00)	nd	0.76 (0.09)	5.50 (0.48)	0.68(0.01)	4.51 (0.08)	1.09(0.18)	3.10 (0.41)	1.64(0.03)	3.05(0.04)
SP8 W	nd	nd	0.84 (0.19)	pu	0.53 (0.28)	2.57 (0.04)	pu	2.60 (0.07)	1.70 (0.04)	3.14(0.06)
SP8 D	0.11 (0.02)	nd	0.74 (0.01)	5.67 (0.21)	0.71 (0.15)	5.62 (010)	1.29 (0.33)	4.98 (0.16)	1.62(0.01)	3.04(0.04)
M 6dS	nd	nd	0.75 (0.14)	nd	0.57 (0.12)	2.57 (0.02)	2.55 (0.58)	2.47 (0.01)	1.70 (0.02)	3.10(0.00)
SP9 D	0.39 (0.02)	nd	0.71 (0.02)	23.64 (0.37)	1.23 (0.03)	5.68(0.16)	8.59 (1.04)	4.13 (0.54)	1.65 (0.02)	3.07(0.05)
SP10 D	0.42 (0.02)	pu	0.73 (0.02)	23.38 (1.37)	1.05(0.01)	6.57 (0.00)	6.35 (0.90)	4.14(0.09)	1.61 (0.01)	3.07(0.01)
W wet season,	D dry season									

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Sample site	PEV	AMP	AMX	PEG	CLX	DCX	NAF	NOR	CIP	ENR
SP1 W	4.07 (0.22)	pu	2.10 (1.19)	1.22 (1.72)	3.27 (0.71)	3.49 (0.01)	3.99 (0.11)	42.93 (3.91)	21.40 (7.57)	29.73 (5.05)
SP1 D	3.34 (0.36)	nd	1.23 (0.01)	nd	2.75(0.14)	3.16 (0.02)	3.66(0.01)	22.40 (0.14)	14.02 (0.15)	19.70 (0.45)
SP2 W	3.82 (0.34)	nd	1.76 (0.08)	$1.59\ (0.07)$	2.96 (0.83)	3.39 (0.05)	3.89 (0.01)	25.31 (0.50)	14.55 (0.90)	19.78 (0.35)
SP2 D	2.90 (0.04)	nd	1.30(0.00)	nd	2.89 (0.27)	3.15 (0.01)	3.62~(0.00)	22.32 (0.06)	14.40 (0.35)	19.32 (0.57)
SP3 W	3.41 (0.43)	nd	1.28 (0.08)	nd	2.81 (0.75)	3.34 (0.03)	3.79 (0.06)	22.41 (0.16)	16.84 (2.44)	25.95 (9.03)
SP3 D	3.14 (0.25)	nd	1.24 (0.07)	nd	2.21 (0.17)	3.17 (0.02)	3.64 (0.01)	22.32 (0.02)	13.94 (0.12)	19.01 (0.15)
SP4 W	4.86 (0.58)	nd	1.40(0.14)	nd	10.37 (0.48)	8.03 (0.17)	3.77 (0.01)	25.58 (2.64)	13.82 (0.11)	21.53 (3.13)
SP4 D	4.08 (1.53)	0.19(0.09)	1.28 (0.03)	3.74 (0.08)	3.42 (0.25)	3.19 (0.04)	3.65 (0.03)	52.19 (2.84)	56.02 (1.16)	20.45 (0.97)
SP5 W	2.99 (0.06)	nd	1.71 (0.57)	nd	3.40 (1.57)	3.30 (0.02)	3.76 (0.06)	22.19 (0.14)	14.08 (0.42)	19.79 (1.04)
SP5 D	3.09 (0.28)	nd	1.39 (0.20)	nd	2.32 (0.36)	3.16 (0.00)	3.63 (0.01)	22.98 (0.97)	14.02 (0.26)	19.76 (0.43)
SP6 W	2.89 (0.01)	nd	1.35 (0.22)	pu	2.14 (0.06)	3.26 (0.00)	3.74 (0.00)	22.16 (0.09)	13.93 (0.21)	19.16 (0.08)
SP6 D	3.09 (0.23)	nd	1.21 (0.01)	0.46(0.03)	2.90 (0.62)	3.15 (0.02)	3.62 (0.00)	22.84 (1.07)	13.92 (0.22)	19.60 (0.27)
SP7 W	3.22 (0.08)	nd	1.30(0.06)	nd	2.47 (0.09)	3.20 (0.03)	3.68 (0.01)	22.48 (0.27)	14.04 (0.08)	21.46 (0.37)
SP7 D	3.03 (0.20)	pu	1.21 (0.01)	nd	2.35 (0.27)	3.14 (0.01)	3.61 (0.01)	27.24 (1.78)	15.40 (1.81)	19.28 (0.44)
SP8 W	3.37 (0.13)	nd	1.47 (0.01)	nd	2.70 (0.37)	3.22 (0.05)	3.70 (0.03)	23.01 (1.11)	13.88 (0.02)	19.67 (0.04)
SP8 D	3.28 (0.36)	nd	1.26 (0.10)	nd	2.82 (0.45)	3.13 (0.00)	3.62 (0.00)	22.38 (0.15)	16.78 (1.76)	21.04 (1.59)
W 6dS	3.17 (0.20)	pu	1.32 (0.09)	nd	2.93 (0.14)	3.18 (0.01)	3.68 (0.01)	22.23 (0.13)	13.68 (0.02)	21.42 (3.25)
SP9 D	3.00 (0.01)	nd	1.35 (0.04)	nd	2.60 (0.75)	3.12 (0.02)	3.61 (0.01)	22.62 (0.45)	13.85 (0.25)	21.97 (2.37)
SP10 D	3.78 (0.33)	nd	1.23 (0.02)	nd	3.08 (0.69)	3.13 (0.01)	3.61 (0.00)	23 33 (1 27)	14 72 (1 08)	19 52 (0 39)

Table 6 Concentrations of MLs, LNs, NIs, TMP and ATs in surface water in ng L^{-1}

Sample site	ERY	TYL	LIN	МЕТ	ТМР	OFX	MEB	ALB
SP1 W	3.53 (0.06)	5.46(0.01)	3.67 (0.00)	3.13 (0.00)	0.50 (0.05)	1.99 (0.05)	2.77 (0.01)	2.89 (0.05)
SP1 D	3.97 (0.00)	5.82 (0.11)	6.43 (0.36)	3.12 (0.01)	1.53 (0.35)	2.21 (0.02)	2.69 (0.00)	2.82 (0.01)
SP2 W	3.47 (0.14)	5.51 (0.12)	3.70 (0.02)	3.12 (0.02)	0.72 (0.09)	2.14 (0.15)	2.74 (0.00)	2.84 (0.02)
SP2 D	3.98 (0.01)	5.46(0.04)	6.69 (0.02)	3.11 (0.00)	0.17 (0.05)	2.02 (0.04)	2.69 (0.01)	2.80 (0.00)
SP3 W	3.32 (0.07)	5.50 (0.08)	4.93 (0.37)	3.12 (0.00)	0.29 (0.13)	2.10 (0.06)	2.71 (0.02)	2.83 (0.01)
SP3 D	3.97 (0.00)	6.63 (0.08)	6.47 (0.71)	3.12 (0.01)	0.66 (0.05)	2.06 (0.02)	2.67 (0.00)	2.82 (0.01)
SP4 W	4.13 (0.05)	8.49 (0.11)	5.66 (0.00)	3.22 (0.12)	2.40 (0.15)	2.30 (0.32)	2.78 (0.01)	4.55 (0.57)
SP4 D	6.73 (0.44)	9.00 (0.76)	6.95 (0.11)	5.56 (0.20)	67.32 (2.87)	2.47 (0.05)	2.71 (0.01)	3.48 (0.03)
SP5 W	4.00 (0.05)	8.83 (0.43)	5.70(0.05)	3.14 (0.01)	0.66 (0.32)	2.09 (0.05)	2.75 (0.04)	2.81 (0.01)
SP5 D	4.01 (0.02)	8.52 (0.06)	7.32 (0.08)	3.15 (0.04)	2.87 (0.00)	1.97 (0.03)	2.67 (0.00)	3.04 (0.00)
SP6 W	4.00 (0.01)	6.96 (0.58)	5.71 (0.02)	3.14 (0.03)	0.44 (0.06)	2.02 (0.00)	2.72 (0.01)	2.81 (0.00)
SP6 D	4.03 (0.07)	8.57 (0.05)	7.63 (0.10)	3.30 (0.27)	1.52 (0.24)	2.11 (0.02)	2.69 (0.02)	3.07 (0.27)
SP7 W	3.98 (0.00)	8.71 (0.37)	6.17 (0.71)	3.14 (0.02)	0.46 (0.20)	2.14 (0.01)	2.70 (0.00)	2.77 (0.01)
SP7 D	4.10 (0.03)	8.85 (0.18)	6.83 (0.22)	3.15 (0.00)	0.64 (0.13)	2.09 (0.04)	2.69 (0.01)	2.95 (0.03)
SP8 W	3.98 (0.02)	8.49 (0.03)	6.32 (0.57)	3.12 (0.01)	0.95 (0.14)	2.12 (0.07)	2.74 (0.01)	2.77 (0.01)
SP8 D	4.05 (0.04)	9.08 (0.34)	7.77 (0.14)	3.17 (0.01)	2.25 (0.75)	2.02 (0.02)	2.70 (0.01)	3.05 (0.06)
SP9 W	3.83 (0.07)	8.49 (0.10)	5.99 (0.00)	3.14 (0.03)	0.30 (0.01)	1.99 (0.04)	2.69 (0.02)	2.80 (0.02)
SP9 D	4.09 (0.10)	8.92 (0.07)	7.68 (0.01)	3.14 (0.00)	1.41 (0.10)	2.06 (0.09)	2.77 (0.01)	5.18 (0.05)
SP10 D	4.13 (0.06)	9.02 (0.05)	7.69 (0.01)	3.25(0.02)	1.49 (0.20)	2.16 (0.06)	3.00 (0.04)	21.39 (7.81)

which increased (average concentration) in the order CIP<ENR<NOR (Tables 5 and S4b). The highest amount of CIP and NOR of 56.02 and 52.19 ng L^{-1} , respectively, were recorded from sampling SP4 during the dry season. The concentrations of the three FQs was variable, and no seasonal trend could be drawn (supplementary Fig. S1b). FQs, like SAs are also commonly found in aquatic systems. In surface waters, they have been reported at varied concentrations, for example, reported CIP concentrations ranged from non-detectable (nd) to 5,015,000 ng L^{-1} in Leça river in Portugal, Yamuna river in India, and Charmoise river and Arc river in France (Dinh et al., 2017; Feitosa-Felizzola & Chiron, 2009; Fernandes et al., 2020; Mutiyar & Mittal, 2014), the highest concentration having been reported in the Ganges river in India (Sharma et al., 2019). Enrofloxacin, ENR, has similarly been studied in surface water with reported concentrations of up to 181,609 ng L^{-1} (Sharma et al., 2019; Yao et al., 2017). Likewise, NOR has reported ranges of 9.57–1261 ng L^{-1} in Leça river, Portugal and Jianghan Plain, Central China (Dinh et al., 2017; Yao et al., 2017), and a high concentration of 16,148-251,137 ng L⁻¹ in Ganges River in India (Sharma et al., 2019). Ofloxacin, though not determined in this study, is also a commonly reported pollutant in surface water at concentrations of up to 11,700 ng L^{-1} (Bhagat et al., 2020). Among the reported concentrations of FQs in surface water in Kenya include NOR at levels of up to 4900 ng L^{-1} and CIP of up to 2800 ng L^{-1} (Kairigo et al., 2020a, b).

Of the two MCs, TLY was detected at higher levels in the range 5.46–9.08 ng $\rm L^{-1}$ and 3.32–6.73 ng $\rm L^{-1}$ for ERY (Tables 6 and S4c). ERY amounts were higher during the dry season (difference not significant) whereas the concentration of TLY varied for the two seasons (supplementary Fig. S1c). ERY has been previously reported in surface water at a range of 32.3–2910 ng L⁻¹ in China (Yao et al., 2017), nd-240 ng L^{-1} in South Africa (Matongo et al., 2015), and TLY at 9.8–74.2 ng L^{-1} in Italy (Zuccato et al., 2005). Concentrations of up to 1.9 μ g L⁻¹ of ERY were reported in Nairobi river water in Kenya (Ngigi et al., 2020). Similar to ERY, the concentration of LIN (lincosamide) ranged from 3.67 to 7.77 ng L^{-1} and was significantly higher during the dry season (supplementary Fig. S1c). There a few studies reporting the occurrence of LIN in surface water, for example, a concentration range of 1.9-248.9 ng L⁻¹ in surface water (rivers, lake, and ponds) (Bhagat et al., 2020; Castiglioni et al., 2005; Feitosa-Felizzola & Chiron, 2009). Amount of MET, a nitroimidazole, was generally low and ranged from 3.11 to 5.56 ng L^{-1} . However, K'oreje et al. (2012) reported a concentration of over 500 ng L^{-1} in their study within Nairobi river basin, Kenya. Concentration of TRM, a diaminopyrimidine, ranged from 0.17 to 67.32 ng L^{-1} , which was higher during the dry season except at sampling point SP2. Notably, is the highest amount of TRM, which was obtained from site SP4 in the dry season. Studies have reported TRM concentrations of up 13,680 ng L^{-1} in surface water (Archundia et al., 2017; Dinh et al., 2017; Fernandes et al., 2020; Zhang et al., 2012, 2020). Concentrations of up to 9480 ng L^{-1} were reported in Kenya for TRM in surface water (K'oreje et al., 2012; Kairigo et al., 2020b; Ngigi et al., 2020) from Nairobi and Mitheu rivers.

Four anthelmintics, ATs (commonly used to treat parasitic worms) from benzimidazole class, were determined in surface water and were obtained in the range 1.97-21.39 ng L⁻¹. FLB was not detected at all sampling points, whereas the other three had a 100% occurrence frequency and their concentrations increased in the order OXF < MEB < ALB (Tables 6 and S4c). The concentrations of MET, OFX, and MEB were similar during the wet and dry season and variable without any obvious tend for ALB. A detection frequency of 77 to 100% was reported for 19 ATs in river water (Chen et al., 2021b), whereas the same study reported a concentration range of nd-61.12 ng L^{-1} for seven BZs, namely, ALB, OXF, MEB, ricobendazole (RIC), fenbendazole (FEN), flubendazole (FLU), and thiabendazole (THI). Other studies have reported up to 197 ng L^{-1} anthelmintics in surface water (Kumirska et al., 2015; Zrnčić et al., 2014); however, very few studies have been done on these pollutants in aquatic environments.

The differences in detected amounts of the antibiotics and benzimidazole anthelmintics in the surface water may be attributed to several factors including physicochemical characteristics of the compounds, usage patterns in the environs, anthropogenic activities, biogeochemical processes, surface runoffs, in addition to environmental fate processes such as biodegradation, photo-degradation, and adsorption (Chen et al., 2018; Kümmerer, 2009; Tang et al., 2015). As a result, the order of detections in different studies may differ. For example, in a study by Duong et al. (2021), the average concentrations (ng L⁻¹) of five classes of antibiotics decreased in the order SAs (117.9) > β -lactams (31.28) > quinolones (20.19) > MLs (17.74) > TMP (8.93). Though, it is notable that the study ranked SAs as the highest and this agrees with the observation in this study. For the detected compounds, there was a negative correlation between the obtained concentrations and log K_{ow} (r = -0.2). However, a positive correlation existed between the concentrations and solubility (r = 0.71) (Table S6a, b in supplementary material).

Largely, all the antibiotics and benzimidazoles determined in this study had concentrations within globally reported values. As a primary source of antibiotics in aquatic systems, final effluents from WWTPs (both hospital and domestic) contain a variety antibiotic residues (Rodriguez-Mozaz et al., 2020) as antibiotics are not fully eliminated (Abuin et al., 2006), which was one of the major contributing factors in the studied river water system, judging from the location. The behavior and fate of antibiotics in the aquatic environment are influenced by the physical and chemical characteristics of the compounds (molecular structure, size, shape, solubility, hydrophobicity, and sorption potentials), environmental factors (pH), and the climatic conditions (temperature and precipitation), as well as biological factors (microbial degradation). Thus, the detection of antibiotics in aquatic systems is greatly affected by their sorption potentials (K_d) in soil particles (among other factors), where surface runoffs and leaching are contributory pathways. Those with low K_d (< 5 L kg⁻¹) and half-life of less than 21 days are easily transported to aquatic systems, whereas those with high K_d (>5 L kg⁻¹) tend to accumulate and persist in the soil matrix. For example, SAs have relatively low K_d of 0.2 to 2.0 L kg⁻¹ and organic carbon normalized partition coefficient $(K_{\rm oc})$, which indicates their low sorption affinity to soil and sediment particles (Thiele-Bruhn, 2003) compared to FQs (K_d of 70 to 5000 L kg⁻¹) (Sarmah et al., 2006; Van Dijk & Keukens, 2000), and are therefore likely to be mobile in the aqueous runoff component following application in soil. However, K_d values vary widely in different types of soils. Moreover, SAs are extensively used to treat human and animal diseases, hence they are ubiquitous in aquatic environments. For SMXZ, it is often used in combination with TRM (Cotrim) because this enhances the effectiveness of the sulfonamide. This combination is moderately mobile and hydrophilic (log K_{ow} values ≤ 0.91), hence

easily transferable into the aquatic environment (Viana et al., 2021), and this may be one of the factors that led to the detection of this compound at higher levels compared to other SAs.

Some antibiotics form conjugated metabolites that may be converted back to parent compounds after excretion. An example is the formation of acetylated metabolites such as N4-acetylsulfamethoxazole or N4-acetylsulfamethazine (in SAs), which can be converted back to the parent form (Viana et al., 2021). Such metabolites have been detected in WWTPs, implying that they form part of residual pollutants that enter recipient surface waters from this source. For most antibiotics, the deconjugation of conjugated active and inactive metabolites into the active parent molecules in WWTPs via biological transformation by microbial enzymes such as glucuronidases and sulfatases, can lead to negative removals of such compounds (example, ERY), hence their discharge into receiving surface waters (Brown & Wong, 2018; Mirzaei et al., 2018).

FQs are considered relatively stable in the environment than other antibiotics, which allows them to persist for longer periods, spreading further and hence accumulating in water and sediments. The fate of these FQs in aquatic systems is mainly dominated by adsorption (log $K_{oc} \ge 4.2$, strong sorption) and photodegradation reactions and therefore rapidly move from water to soil/sediments and onto organic particles solution, hence the occurrence in surface water, although adsorption is more critical (Cardoza et al., 2005; Viana et al., 2021). The fact that FQs are easily adsorbed by sediments, accounts for their detection in low concentrations in surface water as compared to SAs (Li et al., 2021). CIP and ENR are among the most frequently detected FQs. Presence of ENR can contribute to increased amounts of CIP as it is the primary degradation product of enrofloxacin (about 13–60% of ENR is metabolized into ciprofloxacin) (Cardoza et al., 2005; Viana et al., 2021). Lincomycin is moderately mobile and therefore detected in surface water, having been mobilized from primary sources such as domestic and wastewater treatment effluents (Zhu et al., 2020) and from animal manure or animal production (Kuchta & Cessna, 2009).

Benzimidazole anthelmintics are widely used for helminthic infections and are utilized in animal husbandry, agriculture, aquaculture, and human health (Zajíčková et al., 2020). Their detection in the environment, like antibiotics, is through anthropogenic activities, including sewage discharge, agricultural irrigation, and livestock breeding (Sim et al., 2013). BZs were reported to be the dominant class in river water among six other ATs due to their heavy use (Chen et al., 2021b). With respect to the study area, a combination of these sources is possible as the location has several hospitals, residential houses, and some agricultural activities.

Pharmaceuticals in sediments

Sediments were collected at each corresponding water sampling point from Sosiani River and tested for presence of the selected pharmaceuticals using described methods. Residual pharmaceuticals varied for individual compounds in the range of undetected to 26.4 μ g kg⁻¹ (dry weight) for most of the compounds (Fig. 2a-c). However, for the two compounds PEG (PNs) and ALB (BZs), the concentration ranged from 414 to 974 μ g kg⁻¹ and 3–125 μ g kg⁻¹, respectively (Fig. 2d). The frequency of occurrence was 100% for all detected compounds except for two SAs, SDZ and SMZ, with 40 and 20% occurrence, respectively. Residual concentrations in sediments were significantly higher than in surface water (in orders of 10 to 1000 times). Of the eight classes detected in sediments, the amounts followed the decreasing order (based on maximum amounts); PNs>BZs>FQs>MLs>DAPs \approx LNs > NIs > SAs. Out of the 16 SAs analyzed, 8 were detected in the sediments. The quantified amounts for each class ranged as follows: PNs 0.83-973.87, BZs 1.95-124.56, FQs 13.22-26.35, MLs 3.97-11.64, LNs 4.27-6.55, NIs 1.98-5.10, DAPs (TMP) 0.08-6.55, and SAs 0.01–4.13 μ g kg⁻¹. It was also observed that the point SPD 4 had relatively higher amounts compared to other sampling points. Seven compounds were not detected in sediments, which included STZ, SMM, SCP, SFQ, DPS, AMP, and FLB. All pharmaceuticals that were not detected in water were also not detectable in the sediments except for the sulfonamide STZ that was quantifiable in water but not in sediments.

Investigations on occurrence of pharmaceuticals in sediments have been conducted globally and present varied results for antibiotics and other compounds. A report by Chen et al. (2018) placed the average detected amounts of five categories of target antibiotics in sediments in decreasing order, tetracyclines ($88.8 \pm 34.2 \ \mu g \ kg^{-1}$)> fluoroquinolones ($53.7 \pm 33.6 \ \mu g \ kg^{-1}$)> β -lactamases ($33.7 \pm 23.4 \ \mu g \ kg^{-1}$)> others Fig. 2 a Concentrations of SAs in river sediments. b Concentrations of PNs, MET, TRM and OFX in river sediments. c Concentrations of FQs, MLs, and MEB in river sediments. d Concentrations of PEG and ALB in river sediments



 $(11.5 \pm 9.8 \ \mu g \ kg^{-1})$. Of these, SDZ, NOR, CLX, oxcytetracycline, and LIN showed a 100% detection frequency. Among the reported FOs in sediments are CIP, NOR, ENR, and ofloxacin with values of up to 569, 225, 8.19, and 1560 µg kg⁻¹, respectively (Chen & Zhou, 2014; Dinh et al., 2017; Li et al., 2021; Yang et al., 2010). A concentration range from 1700 to 3500 ng g^{-1} dry weight was reported for a number of antibiotics in river sediments, among which were four FQs (CIP, NOR, ofloxacin, and enoxacin) (Dinh et al., 2017). Though SMXZ was not detected in this study, high concentrations of this compound of up to $344 \ \mu g \ kg^{-1}$ have been reported elsewhere (Dinh et al., 2017; Wei et al., 2013). Other reported SAs include SMZ, SPD, SDZ, sulfaquinoxaline, and sulfathiazole, at 19.7, 9.1, 2.07, 0.08–0.9, and 0.6 μ g kg⁻¹, respectively (Chen & Zhou, 2014). The macrolide, ERY, has been detected at a range of $1.5-24.6 \ \mu g \ kg^{-1}$ (Chen & Zhou, 2014) and a range of nd-87.55 μ g kg⁻¹ for TRM in sediments (Matongo et al., 2015). ALB and ricobendazole were reported as the most dominant BZs in sediments, with concentration ranges of nd-596.06 and nd-68.63 ng g^{-1} for the two compounds, respectively (Chen et al., 2021b). Metronidazole was detected at a concentration of up to 1253.5 ng g^{-1} in sediments from a dam in Msunduzi river catchment, South Africa (Matongo et al., 2015).

There are only two studies (to the best of our knowledge) reporting occurrence of antibiotics (up to this moment) in river sediments in Kenya of amounts of up to 474, 26.6, 13.3, 94, 85, and 94 µg kg⁻¹ CIP, NOR, TRM, AMP, SMXZ, and chloramphenicol, respectively (Kairigo et al., 2020b; Kimosop et al., 2016). Environmental concentration of antibiotics in sediments ranges from a few ng kg⁻¹ to few hundreds $\mu g kg^{-1}$ (Chen et al., 2018), hence supporting observed results in this study where the quantified amount ranged from 0.01 to 974 μ g kg⁻¹. Sediments are important sinks for contaminants, hence concentrating antibiotics and other pollutants. Additionally, sediments are in an anaerobic environment that inhibits the degradation of antibiotics and limits exposure to light, hence the concentrations of some antibiotics in sediments are greater than in water (Li et al., 2021). As earlier noted, FQs are easily adsorbed onto sediments (Li et al., 2021), hence their higher concentrations compared to SAs and MLs. Likewise, BZs can be stably adsorbed in sediments under the conditions of suitable pH and appropriate proportion of organic matter (Pavlović et al., 2018).

Comparison of pharmaceutical residues in river surface water and in sediments

Pearson's correlation was used to establish if there was any relationship between concentrations found in surface water and sediments for those compounds that were detected in both compartments. Additionally, pseudopartitioning coefficients $(K_{n,s})$ of the target compounds between water and sediment were also evaluated. In aquatic systems, the sorption is not in equilibrium; hence, $K_{p,s}$ is not always a valuable parameter but is however useful for estimating the sorption capacity of concerned compounds. From Pearson's correlations $(\rho = 0.05)$, concentrations of the compounds in water were positively and significantly correlated to solubility (r=0.71), whereas the concentration in sediments was negatively correlated (not significant) to the solubility (r = -0.02) (Tables S6a, b, supplementary material). Further, the concentrations of the compounds in water had no significant correlation to the concentrations in the sediments (r = -0.09). The $K_{p,s}$ values ranged from 52 to 943, 797 to 322,190, 870 to 919, 1171 to 1262, and 1111 to 7372 L Kg⁻¹ for SAs, PNs, FQs, MLs, and BZs, respectively (Table S5, supplementary material). SAs were among the compounds with low $K_{p,s}$ values, indicating their low adsorption in sediments. $K_{p,s}$ values of over 2000 L kg⁻¹ correspond to highly adsorbed compounds. Hence, the BZs, ALB, and the penicillin PEG were strongly adsorbed in sediments, with potential of accumulation. Other compounds were moderately adsorbed (FQs, MLs, LIN, and MET). However, for such $K_{p,s}$ values, data should be obtained for a considerable period of time for fair representation of sorption equilibrium with relation to sources of the pollutants.

Generally, as observed in this study, concentrations of the determined pharmaceuticals from seven different classes varied in surface water and river sediments. Pharmaceuticals are ubiquitous and are among the major anthropogenic pollutants in the environment. Previous studies have reported presence of antibiotics and other pharmaceuticals in water and river sediments worldwide. Concentrations in surface water range from a few ng L⁻¹ to a few µg L⁻¹ (Bottoni et al., 2010; Chen et al., 2021a; Kuang et al., 2020; Wang et al., 2016). Detected amounts of antibiotics from the classes SAs, FQs, PNs, MLs, β Ls, and TRM in surface water were within reported values (Chen et al., 2018; Kuang et al., 2020; Wang et al., 2016). Though not as widely studied in environment as the antibiotics, ATS have also been reported in surface water and sediments within the same range as antibiotics (Chen et al., 2021b; Kumirska et al., 2015), similar to findings in this study.

Ecological risk analysis

From the RQ_w values, 50% of the compounds (9 out of 18) were of medium ecological risk in aquatic system, whereas 39% presented low risk. These included the PNs (PEV, AMP, and PEG), FQs (NOR and ENR), the macrolides (ERY and TLY), and LIN (Tables 7 and S7-supplementary material). The sulfonamide, sulfamethoxazole (SMXZ), and the fluoroquinolone, ciprofloxacin (CIP), posed a high risk to aquatic life, having RQ_w values of 1.11 and 3.24, respectively. The rest of the evaluated compounds posed no risk. Many pharmaceuticals pose ecological risk because of their continuous usage (some in large consumption) and strong environmental persistence. Previous ecological risk assessment reported SMXZ as having potential to cause medium damage to Daphnia in the aquatic ecosystem (Chen & Zhou, 2014) and was the only SA posing a risk to algae in effluent water, with an risk quotient > 7 in a report by García-Galán et al. (2011). High RQs of 335.5 were reported for sulfachloropyridazine to green algae and 152 to Daphnia magna in ditch water, and the ecological and human health risks caused by sulfonamide mixtures were larger than the individual risks (Qin et al., 2020). In a review by Duan et al. (2022), the sulfonamides SDZ, SMXZ, and SMZ reportedly posed a great risk to the aquatic system. Results by Tang et al. (2015) suggested that SMXZ, ofloxacin, CIP, and ENR in the surface water of Lake Chaohu and inflowing rivers might pose a high risk to algae and plants. Enoxacin, CIP, and SMXZ posed high ecological risks (RQ > 1) to the aquatic organisms Vibrio fischeri, Microcystis aeruginosa, and Synechococcus leopoliensis, respectively, in aquatic environments (Zhang et al., 2012). For the benzimidazole ALB, its occurrence was classified as of medium risk to selected organisms, in the river and water source of Tuojiang River in Sichuan, China (Chen et al., 2021b). It is evident from ecological risk assessments that pharmaceuticals at certain concentrations pose a threat to aquatic environments. Further studies are necessary to fully understand the hazards that these pollutants present, more so considering the human health risks posed by ARB and ARGs present in aquatic environments (Khan et al., 2019).

 Table 7 Risk quotients for the target antibiotics and benzimidazoles in surface water

Antibiotic	Mean (ng L ⁻¹)	Organism	$EC_{50} \ (mg \ L^{-1})$	NOEC ₅₀	AF	PNEC mg L ⁻¹	$\mathbf{RQ}_{\mathbf{w}}$ (mean)	Risk
STZ	7.380	Scenedesmus vacuolatus	13.1	-	1000	0.013	0.001	n
SMZ	10.529	Scenedesmus vacuolatus	19.52	-	1000	0.020	0.001	n
SPD	3.713	Chlorella vulgaris	1	-	1000	0.001	0.004	n
SMXZ	29.919	Synechococcus leopoliensis	0.027	-	1000	0.000	1.108	h
SDMT	1.912	Lemna minor	0.248	-	1000	0.000	0.008	1
PEV	3.477	Microcystis aeruginosa	0.006	-	1000	0.000	0.579	m
AMP	0.190	Microcystis aeruginosa	0.0002	-	1000	0.000	0.950	m
AMX	1.420	Synechococcus leopoliensis	0.00222	-	1000	0.000	0.639	m
PEG	1.753	Microcystis aeruginosa	0.006	-	1000	0.000	0.292	m
NOR	24.596	Vibrio fischeri	-	0.01038	100	0.000	0.237	m
CIP	16.215	Microcystis aeruginosa	0.005	-	1000	0.000	3.243	h
ENR	20.007	Vibrio fischeri	-	0.00288	100	0.000	0.695	m
ERY	4.068	Microcystis aeruginosa	0.023	-	1000	0.000	0.177	m
TYL	7.701	Microcystis aeruginosa	0.034	-	1000	0.000	0.226	m
LIN	6.279	Anabaena Raphidocelis	0.01	-	1000	0.000	0.628	m
MET	3.281	Chlorella Raphidocelis	3.22	-	1000	0.003	0.001	n
OFX	2.108	Vibrio fischeri	2.21	-	1000	0.002	0.001	n
ALB	4.088	Vibrio fischeri	0.77	-	1000	0.001	0.005	n

n no risk, l low risk, m medium risk, h high risk

Conclusion

In this study, 28 antibiotics and 4 benzimidazoles were investigated in surface water and sediments of an urban and suburban river. Their occurrence differed widely according to classes of the pharmaceuticals and was mainly influenced by anthropogenic activities within the sampled location. SAs had the highest contribution in the surface water, and SMXZ had the highest concentration of 247.0 ng L^{-1} . The concentrations in surface water decreased in the order $SAs > TMP > FQs > ATs > PNs \approx MCs \approx LNs > NIs.$ Residual pharmaceuticals in the sediments varied from nd to 974 µg kg⁻¹. PEG and ALB represented the highest concentration in sediments in the range 414–974 µg kg^{-1} and 3–125 µg kg^{-1} , respectively, while the order of detection was PNs>BZs>FQs>MLs>DAPs \approx LNs>NIs>SAs. Ecological risk assessments showed that SMXZ and CIP were of high risk in the surface water, whereas PEV, AMP, PEG NOR, ENR, ERY, TLY, and LIN were of medium ecological risk in the aquatic system. Continuous monitoring of these pollutants is necessary as researcher's endeavor to understand more on their fate in the environment, human, and ecological risks posed by these compounds, development of policies for interventions, and sustainable mitigation strategies.

Author contribution Catherine Chemtai conducted the sampling, did sample preparations, obtained funding from Africa Centre of Excellence in Phytochemicals, Textile, and Renewable Energy (ACE II-PTRE) for her studies, and wrote the initial manuscript draft. Fredrick O. Kengara and Anastasiah N. Ngigi obtained funds, did project conception and design, provided supervision, and contributed to the editing of the manuscript. Anastasiah N. Ngigi was involved in project administration, method validation, and optimization and carried out data analysis and interpretation. All authors reviewed the manuscript.

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Data availability All data for the manuscript is available on request.

Declarations

Conflict of interest The authors declare no conflict of interest to declare that are relevant to the content of this article.

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