The availability, price and affordability of essential antibacterials in Hubei province, China

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Abstract
Background: China ranks first amongst the countries for the abuse of antibacterials. Essential antibacterials could help solve the problem. The aim of the work is to evaluate the availability, price and affordability of essential antibacterials in Hubei province, China.

Method: The standardized methodology developed by the World Health Organization and Health Action International was used to collect data on the availability and prices of 16 antibacterials in 5 cities of Hubei province, China.

Results: First, in total, the median availability of originator brands and lowest-priced generics for the essential antibacterials was low, 3.0% (0.0, 18.2%) and 33.3% (0.0, 87.9%) for each, respectively. Second, the median price ratio of originator brands for the antibacterials was 20.30 (4.71, 35.80), while for generics, it was 0.49 (0.07, 1.18). Third, the affordability of originator brands for the antibacterials was 0.35 (0.04, 6.11). Finally, we found that in Hubei province, lowest-priced generics for essential antibacterials with (fairly) high availability and relatively low price included Amoxicillin/Clavulanic Acid, Cefazidime, Metronidazole, Gentamicin Sulfate and Ceftriaxone.

Conclusion: The prices of lowest-priced generics for essential antibacterials in Hubei province were reasonable, and in tertiary hospitals the availability was the highest, while in secondary and primary hospitals, it was relatively lower. Originator brands were not only extremely expensive but also difficult to obtain. Measures should be taken to improve the availability of essential antibacterials and the affordability of originator brands.

Keywords: Essential antibacterials, Availability, Price, Affordability, China

Background
Antibacterials are medicines that are used to prevent and treat bacterial infections [1, 2]. Ever since penicillin was introduced into medical therapy, hundreds of antibacterials have been isolated or synthesized to save lives from infectious diseases [3]. As more and more antibacterials are provided, overuse and misuse of them have been a serious problem. The overuse and misuse of antibacterials stimulated more rapid growth of antibacterial-resistance bacteria, reducing their therapeutic potential against human and animal pathogens [4]. World Health Organization (WHO) characterises antimicrobial resistance as a global public health crisis which must be managed with the utmost urgency [5]. According to the WHO, overuse of antibacterials will cause 10 million deaths and a loss of 100,000 billion US dollars by 2050 [6].

Overusing and misusing antibacterials is a key driver of bacteria resistance. When a new antibacterial is introduced into medical usage, drug resistance to it follows naturally, but it sometimes comes swiftly [7]. Overusing antibacterials concerns more about consumption. China had the largest consumer group of antibacterials besides Brazil, Russia, India and South Africa [8]. In 2010, China was the second largest consumption of antibacterials with 10.0 × 10^9 units, following India [8]. While in 2013, 92,700 t of antibacterials were consumed in China, and 48% of which were used in humans [3]. From
2000 to 2010, consumption of antibiotics in hospitals among developing countries had a great increase, in which 57% was attributed to China [8]. As estimated, each Chinese person receives treatment by antibiotics in an average amount of 138 g a year, which is 10 times that consumed in the United States [9]. Further, the rate of antibacterial prescriptions for inpatients is 80%, compare with just 10% in hospitals in high-income countries, while the WHO recommends a maximum of 30% [10, 11]. Problem of antibiotics misuse in China comes as using too many high-level antibacterials instead of essential antibacterials. In Hubei province, essential antibacterials were not fully used, while Cefotaxime, Cefoperazone, Cefamandole, Levofloxacin, Amikacin are preferred (Additional file 1) [12–14]. Similarly, in the newest report, quinolone, 3rd generation cephalosporings, cephalosporings with enzyme inhibitor, 2nd generation cephalosporings, and penicillins with enzyme inhibitor were the 5 kinds of antibacterials that were used most frequently in public hospitals [15].

Owing to the overuse and misuse of antibiotics, China has a rapid growth rate of antibacterial resistance. A high prevalence of antibacterial resistance occurs among healthcare-associated pathogens such as mycobacterium tuberculosis, Gram-negative organisms and Gram-positive organisms [16]. Most antibacterials except amoxicillin/clavulanic acid, levofloxacin and chloramphenicol, have high resistance [17]. These facts caused worldwide concern. Scientists said that in the foreseeable future, carbapenem resistance in China would be serious [18]. In China, many researchers have focused on the problem of antibacterial overuse and misuse (Additional file 2). Studies showed that even the well-educated groups in China have problems with antibacterial use, we can infer that the situation across China is no more optimistic. The reason for this could be concluded as patients’ expectations, financial incentives from pharmaceutical suppliers, hospital income from drug sale profits and fragmentary, incomplete national guidance on the use of antibacterials that led to antibacterial overuse [19].

Misuse or overuse of antibacterials could lead to severe resistance and loss of money, and choosing effective essential drugs could help reduce overuse of antibacterials and slow down antibacterial resistance [20]. For this purpose, the policy of having a national essential medicine list was rolled out and could help restrain the problem from the source. The policy requires patients to get rational and efficient treatment [21]. Availability and affordability are the two main criteria to assess whether patients can receive timely, adequate and efficient treatment [22]. Until now, several surveys have been conducted to study the availability and affordability of (essential) medicines in China to evaluate the progress of the new policy [21, 23–25]. However, to our knowledge, so far, there has not been a survey to study the availability and affordability of antibacterials alone in Hubei province, so this study aims to determine the high-availability, low-cost antibacterials for three levels of hospitals in Hubei province, based on the most recently published 19th WHO Model List of Essential Medicines to make a suggestion regarding the fight against antibacterial resistance.

**Objective**

This study aimed to evaluate the effects of essential medicine policy on the availability of antibacterials in Hubei province and to assess the prices of originator brands (OBs) and lowest-priced generics (LPGs) for essential antibacterials. This study was also conducted to find essential antibacterials with high availability and low price for three levels of hospitals in Hubei province.

**Method**

This study was conducted in 33 hospitals from 5 cities, including Wuhan, Yichang, Xiangyang, Jingzhou and Huanggang (Table 1), in Hubei province, central China, from August to October in 2016, using a questionnaire mainly containing the surveyed information of drugs’ common name, dosage, brand name, supplied dosage, price etc. which is developed by the WHO/HAI. Survey data were entered into the pre-programmed Excel Workbook (WHO/HAI 2008) by two people using a double entry technique. Sixteen essential antibacterials were included based on the 19th WHO Model List of Essential Medicines and Essential Medicines List for Hubei province in 2014 (Additional file 3). For each medicine, data on the availability and price of OBs and LPGs were collected. Prices were recorded on the day of the survey. The unit prices were calculated and entered into the workbook.

The availability of each medicine was reported as the percentage of outlets in which the surveyed medicine could be offered on the day of data collection. The median price ratio (MPR), or the ratio of one medicine’s median unit price to the international reference prices (IRPs) according to the Management Sciences for Health

**Table 1 Samples of facilities**

| Districts  | Primary | Secondary | Tertiary | Total |
|-----------|---------|-----------|----------|-------|
| Wu Han    | 2       | 2         | 2        | 6     |
| Yi Chang  | 2       | 2         | 2        | 6     |
| Xiang Yang| 2       | 3         | 2        | 7     |
| Jing Zhou | 3       | 2         | 2        | 7     |
| Huang Gang| 2       | 4         | 1        | 7     |
| Total     | 11      | 13        | 9        | 33    |
Price Indicator Guide for 2015, was used for price evaluation. To assess affordability, the ratio of the medicine prices in a standard treatment to the average daily wage of the lowest-paid unskilled government workers could be used. Due to the lack of official data on daily wages of the lowest-paid workers, we used the mean minimum wage in Hubei province instead of it, which was RMB 36.67 (USD 5.50) at the survey time.

Availability, price and affordability: Criteria in the study

Availability

- Absent, 0% of hospitals: the medicines were not found in any surveyed hospitals.
- Low, < 50% of hospitals: the medicines were hard to be found.
- Fairly high, 50–80% of hospitals: the medicines were available in many hospitals.
- High, > 80% of hospitals: the medicines have good availability.

Price

The MPR for each medicine was calculated using the Workbook developed by WHO/HAI, only if the medicine was available in at least one facility. The MPR is the local median unit price of a medicine in comparison with the median unit price found in the Management Sciences for Health (MSH) Price Indicator Guide for 2015 (MSH 2015). The ideal value for MPR was used to represent acceptable local price ratios:

- Acceptable line: MPR < 1;
- Warning line for generics: MPR > 2

According to the WHO/HAI methodology, if an MPR is twice the international reference price for a generic equivalent product (considered a warning line) [26], then this should be a cause for concern.

Affordability

Affordability was estimated by comparing the total cost of a medicine for a standard course of treatment to the daily wages of the lowest paid unskilled government worker, which was 36.67 CNY per day at the time of the survey.

- Good: Affordability < 1

Results

Availability

Generally, both OBs and LPGs of the antibacterials had low availability in the surveyed hospitals, with median availability 3.0% (0.0, 18.2%) for OBs and 33.3% (0.0, 87.9%) for LPGs (Table 2). Among the OBs, Ceftriaxone, Ceftazidime, Ciprofloxacin, Cefaclor and Linezolid could be offered at the study time, though the availability was low, while Amoxicillin/Clavulanic Acid, Ampicillin, Cephalexin and Roxithromycin were absent. Differently, for LPGs, Amoxicillin/Clavulanic Acid and Gentamicin Sulfate had high availability (84.8 and 87.9%), while Linezolid was absent because there were no generic equivalents in China (Table 2 and Table 3). Table 2 shows that the median availabilities of LPGs in primary and secondary hospitals were nearly the same, 36.4% (0.0, 100%) for primary hospitals and 34.7% (0.0, 84.6%) for secondary hospitals. The LPGs with high availability in primary hospitals included Amoxicillin/Clavulanic Acid (100%), Ceftriaxone (90.9%), and Gentamicin Sulfate (90.9%), while Chloramphenicol was completely unavailable. In the secondary hospitals, Ceftazidime, Norfloxacin and Metronidazole had high availability (all 84.6%), while drugs with high availability in the primary hospitals were all in the fairly high-level category. Likewise, LPG of Chloramphenicol was absent in the secondary hospitals. For tertiary hospitals, several OBs of antibacterials could be offered, including Ceftriaxone, Ceftazidime, Ciprofloxacin, and Cefaclor as well as Linezolid, and the median availability was low (11.1%). For LPGs, Ceftriaxone (88.9%), Ceftazidime (88.9%), Gentamicin Sulfate (100.0%), Norfloxacin (88.9%), and Metronidazole (100.0%) had high availability. In addition, the only antibacterial without an LPG was Linezolid, while Chloramphenicol had 11.1% availability.

Medicine prices

As shown in Table 4, in the surveyed hospitals, the median MPRs for 3 OBs and 11 LPGs were 20.30 (4.71, 35.8) and 0.49 (0.07, 1.18) times their IRPs, respectively. For 3 medicines available as both OBs and LPGs, including Ceftriaxone, Ceftazidime and Ciprofloxacin, OBs cost nearly 30 times more than LPGs. The OB of Ciprofloxacin was extremely expensive in the surveyed hospitals, such that the price was 35.8 times its IRP. All LPGs, except Amoxicillin/Clavulanic Acid and Ciprofloxacin, had lower prices than the IRP. In fact, the MPRs of Amoxicillin/Clavulanic Acid and Ciprofloxacin were both 1.18, below the warning line (MPR = 2.0). The lowest MPR of OBs belonged to Nitrofurantoin (0.07).

Affordability

Table 5 shows the affordability of treatments with 3 OBs and 11 LPGs. All OBs cost more than 1 day's wages. The median affordability of OBs was 28.14 (21.70, 41.90). In addition, the generic equivalents of the OBs all cost less, with 0.46 for Ceftriaxone (vs. 21.70), 6.11 for Ceftazidime (vs. 41.90), and 0.93 for Ciprofloxacin (vs. 28.14). Generally, choosing LPGs for treatment cost less. The median affordability of LPGs was 0.35 (0.04, 6.11). In
fact, only Ceftazidime and Piperacillin/Tazobactam cost more than 1 day's wages, with affordability of 6.11 and 2.58, respectively, for each.

**Comprehensive analysis of medicine availability and price**

Fig. 1 depicts the availability and prices of both OBs and LPGs for the surveyed antibacterials. For medicines located in the 2nd quadrant, patients had good access and relatively low prices; for example, the LPG for Gentamicin Sulfate had 87.9% availability and an MPR of 0.49. Generics located in the 1st quadrant were medicines with high availability, but with prices higher than the IRPs, including Amoxicillin/Clavulanic Acid. For generics in the 3rd quadrant, the prices were low, but patients had difficulty getting them in the hospitals; for example, though the MPR of Chloramphenicol was 0.40, it only had 3.0% availability. For generics in the 4th quadrant, patients not only had to pay higher prices than the reference prices but were also not able to easily obtain the medicines; for example, the LPG for Ciprofloxacin had 18.2% availability and an MPR of 1.18. In contrast, all LPGs of the surveyed essential antibacterials in Hubei province had MPR values below the warning line (MPR = 2.0). For OBs, the three antibacterials were all located in the 4th quadrant, including Ceftazidime, Ceftriaxone and Ciprofloxacin.

Overall, LPGs of essential antibacterials had reasonable prices, and the availability ranged widely, while OBs were not only difficult to obtain but also expensive.

**Discussion**

There are four main findings in this essay. First, in total, the median availabilities of OBs and LPGs for the essential antibacterials were low. The low availability of essential antibacterials could be explained by the following reasons. First, the government's compensation mechanism was not clear, so hospitals had no willingness to use essential medicines, which brought less profit. Second, the public had the opinion that essential medicines were cheap and old drugs, which were less powerful than expensive and new drugs, so hospitals and doctors were pressured against using essential medicines. Third, the research and development ability of pharmaceutical companies grew rapidly, more and more new drugs were advertised, and essential drugs lacked propaganda; as a result, doctors tended to use new drugs with advertised better effects and less adverse actions. Finally, a handful of doctors might be impelled by economic benefits not to prescribe essential medicines for patients, so essential medicines were likely to be eliminated.

Second, the median MPR of OBs for the antibacterials was 20.30 (4.71, 35.80), while for LPGs, it was 0.49 (0.07, 1.18), which was at an acceptable level. In fact, all LPGs for the antibacterials could be acquired with acceptable prices. It was clear that OBs for the surveyed antibacterials were extremely expensive, while prices of generic equivalents were quite acceptable. In fact, in China, the cost of OBs should not exceed LPGs by 35% or more [21], but in our survey, the OB cost at least 6.8 times

| Number | Medicine                    | LEML  | Total Availability | Availability in Primary Hospitals | Availability in Secondary Hospitals | Availability in Tertiary Hospitals |
|--------|-----------------------------|-------|--------------------|-----------------------------------|------------------------------------|-----------------------------------|
|        |                             |       | OBS     | LPGs   | OBS     | LPGs   | OBS     | LPGs   |
| 1      | Amoxicillin/Clavulanic Acid | Y     | 0.0%    | 84.9%  | 0.0%    | 100.0% | 0.0%    | 76.9%  | 0.0%    | 77.8%  |
| 2      | Ampicillin                  | Y     | 0.0%    | 24.2%  | 0.0%    | 45.5%  | 0.0%    | 15.4%  | 0.0%    | 11.1%  |
| 3      | Ceftriaxone                 | Y     | 12.1%   | 75.8%  | 0.0%    | 90.9%  | 0.0%    | 7.7%   | 53.8%   | 33.3%  |
| 4      | Ceftazidime                 | Y     | 3.0%    | 69.7%  | 0.0%    | 36.4%  | 0.0%    | 84.6%  | 11.1%   | 88.9%  |
| 5      | Chloramphenicol             | N     | –       | 3.0%   | –       | 0.0%   | –       | 0.0%   | –       | 11.1%  |
| 6      | Ciprofloxacin               | Y     | 3.0%    | 18.2%  | 0.0%    | 18.2%  | 0.0%    | 15.4%  | 11.1%   | 22.2%  |
| 7      | Nitrofurantoin              | N     | –       | 30.3%  | –       | 27.3%  | –       | 30.8%  | –       | 33.3%  |
| 8      | Cephalexin                  | Y     | 0.0%    | 33.3%  | 0.0%    | 18.2%  | 0.0%    | 30.8%  | 0.0%    | 55.6%  |
| 9      | Gentamicin Sulfate          | N     | –       | 87.9%  | –       | 90.9%  | –       | 76.9%  | –       | 100.0% |
| 10     | Norfloxacin                 | N     | –       | 69.7%  | –       | 36.4%  | –       | 84.6%  | –       | 88.9%  |
| 11     | Metronidazole               | Y     | –       | 78.8%  | –       | 54.5%  | –       | 84.6%  | –       | 100.0% |
| 12     | Piperacillin+Tazobactam     | Y     | –       | 21.2%  | –       | 18.2%  | –       | 7.7%   | –       | 44.4%  |
| 13     | Cefaclor                    | Y     | 18.2%   | 33.3%  | 18.2%   | 45.5%  | 0.0%    | 38.5%  | 44.4%   | 11.1%  |
| 14     | Roxithromycin               | Y     | 0.0%    | 60.6%  | 0.0%    | 63.6%  | 0.0%    | 53.8%  | 0.0%    | 66.7%  |
| 15     | Furazolidone                | N     | –       | 27.3%  | –       | 18.2%  | –       | 23.1%  | –       | 44.4%  |
| 16     | Linezolid                   | N     | 3.0%    | –       | 0.0%    | –       | 0.0%    | –       | 11.1%   | –      |
| Median |                             |       | 3.0%    | 33.3%  | –       | 36.4%  | –       | 34.7%  | 11.1%   | 50.0%  |

LEML Local essential medicine list, OBs Originator brands, LPGs Lowest-priced generics
more than the LPG for Ceftazidime. The high prices of OBs might be due to the bad profit mechanisms when OBs were imported into China. Prices could be elevated several times when crossing levels of related organizations. Luckily, prices for all generics in Hubei province were below the warning line (MPR = 2.0), and only Amoxicillin and Ciprofloxacin had MPRs > 1. Reported by Hao Yang et al. in 2009 [23], MPRs of Amoxicillin and Ciprofloxacin were under 1.5, which was similar to our research. In the survey previously conducted in Shaanxi province [24], which also focused on essential medicines, the authors found that the procured prices of 28 generics were 0.75 times their reference prices. This finding reveals that the prices of generics for essential medicines were low and reasonable. It was not surprising that LPGs for essential antibacterials were cheap because the cost of generics in research and development was omitted. Thus, government should intensify price negotiations with pharmaceutical companies to decrease OB prices and take measures to supervise the abnormal price increases.

**Table 3** Availability of essential antibacterials in the surveyed hospitals

| Availability Range | Total OBs | Total LPGs | Primary Hospitals | Secondary Hospitals | Tertiary Hospitals |
|--------------------|----------|------------|-------------------|---------------------|-------------------|
| Absent 0% | Amoxicillin/Clavulanic Acid | Linezolid | The rest | Chloramphenicol | The rest |
| Low < 50% | Ceftriaxone | Ampicillin | Cefaclor | Ampicillin | Ceftriaxone |
| | Ceftriaxone | Chloramphenicol | Ceftriaxone | Clavulanic Acid | Ceftriaxone |
| | Ciprofloxacin | Ceftriaxone | Ciprofloxacin | Ciprofloxacin |
| | Cefaclor | Nitrofurantoin | Cefaclor | Nitrofurantoin |
| | Linezolid | Cephalixin | Linezolid | Cephalixin |
| | Piperacillin+Tazobactam | Norfloxacin | Piperacillin+Tazobactam | Piperacillin+Tazobactam |
| | Cefaclor | Furazolidone | Cefaclor | Furazolidone |
| Fairly High 50–80% | None | Ceftriaxone | None | Metronidazole |
| | Ceftriaxone | None | None | None |
| | Norfloxacin | None | None | None |
| | Metronidazole | None | None | None |
| | Roxithromycin | Clavulanic Acid | Roxithromycin |
| | Gentamicin Sulfate | Clavulanic Acid |
| | High > 80% | None | None | None |
| | None | None | None | None |
| | Amoxicillin/Clavulanic Acid | Clavulanic Acid |
| | Gentamicin Sulfate | Gentamicin Sulfate |

**Table 4** MPR of 11 kinds of antibacterials in the surveyed hospitals

| No | Medicine | MPR OBs | MPR LPGs |
|----|----------|---------|---------|
| 1  | Amoxicillin/Clavulanic Acid | – | 1.18 |
| 2  | Ampicillin | – | 0.16 |
| 3  | Ceftriaxone | 20.30 | 0.43 |
| 4  | Ceftazidime | 4.71 | 0.69 |
| 5  | Chloramphenicol | – | 0.40 |
| 6  | Ciprofloxacin | 35.8 | 1.18 |
| 7  | Nitrofurantoin | – | 0.07 |
| 8  | Cephalixin | – | 0.83 |
| 9  | Gentamicin Sulfate | – | 0.49 |
| 10 | Metronidazole | – | 0.67 |
| 11 | Piperacillin+Tazobactam | – | 0.17 |
| Median | 20.30 | 0.49 |

*MPR Median price ratio, OBs Originator brands, LPGs Lowest-priced generics*
Third, the affordability of OBs for the antibacterials was 28.14 (21.70, 41.90) times the daily wage of an unskilled government worker, while for LPGs, the affordability was 0.35 (0.04, 6.11) times the daily wage. Due to the high price of OBs, the affordability of OBs was poor. Lowering the prices of OBs is extremely urgent. On the other hand, in general, the affordability of LPGs was quite reasonable. All but Ceftazidime and Piperacillin/Tazobactam had affordability below 1. According to Table 2, prices of LPGs for these two antibacterials were less than their international reference prices, so the prices were acceptable. Hence, for Ceftazidime and Piperacillin/Tazobactam, government should increase the insurance and payment reimbursement ratio to improve the affordability of these two medicines.

Finally, we found that in Hubei province, LPGs for the essential antibacterials with (fairly) high availability and relatively low price included Amoxicillin/Clavulanic Acid, Ceftazidime, Metronidazole, Gentamicin Sulfate and Ceftriaxone. These essential medicines could cover most

### Table 5
Affordability of 11 kinds of antibacterials in the surveyed hospitals

| Number | Description                  | Strength     | DDD   | Duration Days | Affordability |
|--------|------------------------------|--------------|-------|---------------|---------------|
| 1      | Amoxicillin/Clavulanic Acid  | 500 mg/125 mg| 1 g   | 7             | –             | 0.35          |
| 2      | Ampicillin                   | 500 mg       | 2 g   | 7             | –             | 0.31          |
| 3      | Ceftriaxone                  | 1 g          | 2 g   | 7             | 21.70         | 0.46          |
| 4      | Ceftazidime                  | 1 g          | 4 g   | 7             | 41.90         | 6.11          |
| 5      | Chloramphenicol              | 250 mg       | 3 g   | 7             | –             | 0.05          |
| 6      | Ciprofloxacin                | 2 mg/ml      | 0.5 g | 7             | 28.14         | 0.93          |
| 7      | Nitrofurantoin               | 100 mg       | 0.2 g | 7             | –             | 0.04          |
| 8      | Cephalexin                   | 250 mg       | 2 g   | 7             | –             | 0.40          |
| 9      | Gentamicin Sulfate           | 40 mg/ml     | 0.24 g| 7             | –             | 0.23          |
| 10     | Metronidazole                | 200~250 mg   | 1.5 g | 7             | –             | 0.04          |
| 11     | Piperacillin+Tazobactam      | 4 g + 500 mg | 14 g  | 7             | –             | 2.58          |

**Median** 28.14 0.35

**DDD** Defined daily dose, **OBs** Originator brands, **LPGs** Lowest-priced generics
pathogens in common infections. We suggest that doctors and decision-makers should consider using these essential antibacterials more.

**Conclusion**

Chinese NEML policy requires every public hospital to stock essential medicines, including antibacterials, but the availability in Hubei province was still low. The high prices and poor affordability of originator brands might influence their use. Optimistically, the lowest-priced generics might make up for this situation, but more things still need to be done. Choosing essential antibacterials is a way to promote rational drug use because it excludes profit factors, and using the appropriate essential antibacterials could help decrease high-level antibacterial overuse with the purpose of reducing antibacterial resistance.

**Additional files**

**Additional file 1:** The list of common antibacterials. Representative examples of common antibacterials classified by their therapeutic class and the most frequently used kinds in Hubei province. (DOCX 16 kb)

**Additional file 2:** Articles about misuse of antibacterials in China. A brief review of articles about misuse of antibacterials in China, according to the published years [26–31]. (DOCX 16 kb)

**Additional file 3:** The list of surveyed antibacterials. List of the surveyed medicines including their therapeutic class, name, type, dosage form, and belonged catalogues. (DOCX 16 kb)

**Additional file 4:** International Ethical Guidelines for Biomedical Research Involving Human Subjects. A guideline of how ethics committee censors research on human subjects. (PDF 166 kb)

**Abbreviations**

HAI: Health Action International; IRP: International Reference Price; LPGs: Lowest-Priced Generics; MPR: Median Price Ratio; NEML: National Essential Medicine List; OBs: Originator Brands; R&D: Research and Development; WHO: World Health Organization

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**Availability of data and materials**

All data generated or analysed during this study are included in this published article.

**Authors’ contributions**

GSW and DYF performed the study design and revised the manuscript. CHB made substantial contributions to the collection of data. WGW analysed and interpreted the data, as well as drafting the manuscript. All authors read and approved the final manuscript.

**Competing interest**

The authors declare that they have no competing interests.

**Ethics approval and consent to participate**

All the selected hospitals agreed to participate in the study, and a written informed consent form was signed by E-mail before data collection which was approved by the group involved in ethics or other permissions in each surveyed hospital. The ethics committee of the University of Science and Technology (HUST) ruled that no formal ethics approval was required in this case, as no human subjects or other animals were involved in the study complying with the International Ethical Guidelines for Biomedical Research Involving Human Subjects (Additional file 4), as well as the open policy for data of medicine prices.

**Consent for publication**

Not applicable.

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**References**

1. Zaman A, Razzaque S. Designing novel Antibacterials: application of omics science. Klimk Dergisi. 2013;26(1):28–9.
2. Zhang Y. Antibiotics: from modern medicine to global risk; 2017.
3. Qiao M, Ying GG, Singer AC, et al. Review of antibiotic resistance in China and its environment. Environ Int. 2018;110:160–72.
4. Wright GD. Antibiotic resistance in the environment: a link to the clinic? Curr Opin Microbiol. 2010;13:589–94.
5. WHO. Global Action Plan on Antimicrobial Resistance. 2015
6. http://en.people.cn/n3/2016/0622/c90000-9061399.html. Accessed 22 May 2016.
7. Shallcross LJ, Davies DSC. Antibiotic overuse: a key driver of antimicrobial resistance. Brit J Gen Pract. 2014;64:604.
8. Van Boeckel TP, S G A, et al. Global antibiotic consumption 2000 to 2010: an analysis of national pharmaceutical sales data. Lancet Infect Dis. 2014;14:742–50.
9. Li Y. China’s misuse of antibiotics should be curbed. 8mj. 2014:348g1083.
10. Ross-Degnan D, et al., The Development of Standard Indicator Values for the WHO Drug Use Prescribing Indicators. 2013. http://archives.who.int/icium/icium1997/posters/ia2_bx.html.
11. WHO. Medicines use in primary care in developing and transitional countries. 2009. http://www.who.int/entity/medicines/publications/primary_care_8April09.pdf. Accessed 8 Apr 2009.
12. Wang C, Zhang WZ, Song T. Utilization of antibiotics in our hospital in 2009. China Med Herald. 2010;07(22):209–10.
13. Du JM HT. Application and analysis of antibiotics in hospitals. Chin J Aesth Med. 2012;21(18):169.
14. Zhou R. Application of antibiotics in outpatient dispensary of our hospital. Med Front. 2018;8(2):373–4.
15. http://baijiahao.baidu.com/s?id=1616897733034178312&wfr=spider&for=pc. Accessed 12 Nov 2018.
16. Yezi S, Li H. Antibiotic resistance amongst healthcare-associated pathogens in China. Int J Antimicrob Ag. 2012;40:389–97.
17. Hu F, Zhu D, Wang F, et al. Results from the survey of antibiotic resistance (SOAR) 2009-11 and 2013-14 in China. J Antimicrob Chemother. 2016;71(Suppl 1):i33.
18. Paterson DL, Van DD. China’s antibiotic resistance problems. Lancet Infect Dis. 2017;17:351.
19. Reynolds L, Mckee M. Factors influencing antibiotic prescribing in China: an exploratory analysis. Health Policy. 2009;90:32–6.
20. Xu W, Yao X, et al. Clinical study of bacterial resistance and antibiotic use intensity in nosocomial infection. China Pract Med. 2017:12:20–2.
21. Jiang M, Zhou Z, Wu L, et al. Medicine prices, availability, and affordability in the Shaanxi Province in China: implications for the future. Int J Clin Pharm Net. 2015;37:12.
22. Gong S, Wang Y, Pan X, et al. The availability and affordability of orphan drugs for rare diseases in China. Orphanet J Rare Dis. 2016;11:1–12.
23. Yang H, Dib HH, Zhu M, et al. Prices, availability and affordability of essential medicines in rural areas of Hubei Province, China. Health Policy Planning. 2010;25:219.
24. Jiang M, Yang S, Yan K, et al. Measuring access to medicines: a survey of prices, availability and affordability in Shaanxi province of China. PLoS One. 2013;8:e70836.
25. Xi X, Li W, Li J, et al. A survey of the availability, prices and affordability of essential medicines in Jiangsu Province, China. BMC Health Serv Res. 2015;15:345.
26. Lv B, Zhou Z, Yu G, et al. Knowledge, attitudes and practices concerning self-medication with antibiotics among university students in western China. Tropical Med Int Health. 2014;19:769–79.
27. Zhu X, Pan H, Yang Z, et al. Self-medication practices with antibiotics among Chinese university students. Public Health. 2015;130:78–83.
28. Wang XM, Zhou XD, Hesketh T. Massive misuse of antibiotics by university students in China: a cross-sectional study. LANCET. 2016;388:594.
29. Wang X, Peng D, Wang W, et al. Massive misuse of antibiotics by university students in all regions of China: implications for national policy. Int J Antimicrob Ag. 2017;50(3):441–6.
30. Chang J, Ye D, Lv B, et al. Sale of antibiotics without a prescription at community pharmacies in urban China: a multicentre cross-sectional survey. J Antimicrob Chemother. 2017;72:1235.
31. Hu Y, Wang X, Tucker JD, et al. Antibiotic use among future health professionals: A multicentre cross-sectional study of Chinese medical student. Open Forum Infect Dis. 2017;4(Suppl 1):S257.