

Supplementary data:

Table 1: Search strategy

| No. | Searches | Results | | |
|-----|---|---------|---------|---|
| | | Embase | Medline | International Pharmaceutical Abstract |
| 1 | Counterfeit* | 477 | 296 | 301 |
| 2 | Fake | 631 | 491 | 22 |
| 3 | Substandard | 1017 | 874 | 78 |
| 4 | Falsified | 211 | 182 | 10 |
| 5 | 1 or 2 or 3 or 4 | 2230 | 1765 | 375 |
| 6 | Drug* | 2942573 | 1240863 | 248285 |
| 7 | Medicine* | 517065 | 379325 | 23766 |
| 8 | Pharmaceutical* | 62754 | 74697 | 47666 |
| 9 | Antimicrobial* | 61758 | 48954 | 7876 |
| 10 | Antimalaria* | 16651 | 14579 | 3147 |
| 11 | Antibiotic* | 311391 | 146476 | 24572 |
| 12 | 6 or 7 or 8 or 9 or 10 or 11 | 3520198 | 1708464 | 302874 |
| 13 | 5 and 12 | 833 | 522 | 346 |

Table 2: Categories of different issues of tested medicines

| Stated problem | Description |
|--|--|
| Content assay of active ingredient: Inadequate active ingredient excessive active ingredient No active Ingredient | Quantification of the active ingredient content of a drug with regard to claim content declared on the packaging; the result should be within the specified range. |
| Wrong active ingredient | Detection of active ingredient in the drug that is not declared on the packaging |
| Dissolution failure | Solubility or release of active ingredients is not within the specified time range. |
| Presence of impurity | Coexistence of a substance with a drug, such as starting material, intermediates or that is formed as a result of any side reactions. |
| Fake packaging | Packaging has mislabelling information about a drug origin or authenticity |
| Mass uniformity test failure | The weight of a tablet or capsule is not within the average range specified |
| Unknown ingredient | Extraneous contaminants that should not present in a drug |

Table 3: Studies excluded after applying quality assessment criteria

| No. | Studies | Methodological strength scoring (0-12) |
|-----|---|--|
| 1 | Stenson B, et al. The quality of drugs in private pharmacies in the Lao People's Democratic Republic. <i>Int J Risk Saf Med</i> 1998; 11 (4):243-9. | 5 |
| 2 | Atemnkeng MA, et al. Quality control of active ingredients in artemisinin-derivative antimalarials within Kenya and DR Congo. <i>Trop Med Int Health</i> 2007; 12 (1):68-74 | 5 |
| 3 | Ogwal-Okeng J, et al. Quality of oral and parenteral chloroquine in Kampala. <i>East Afr Med J</i> 1998; 75 (12):692-4 | 5 |
| 4 | Ofori-Kwakye K, et al. Quality of Artesunate Tablets Sold in Pharmacies in Kumasi, Ghana. <i>Trop J Pharm Res</i> 2008; 7 (4):1179-84 | 5 |
| 5 | Minzi OMS, et al. Evaluation of the quality of amodiaquine and sulphadoxine/pyrimethamine tablets sold by private wholesale pharmacies in Dar Es Salaam Tanzania. <i>J Clin Pharm Ther</i> 2003; 28 (2):117-22 | 5 |
| 6 | Tipke M, et al. Substandard anti-malarial drugs in Burkina Faso. <i>Malar J</i> 2008; 7 (1):95 | 5 |
| 7 | Newton PN, et al. A Collaborative Epidemiological Investigation into the Criminal Fake Artesunate Trade in South East Asia. <i>PLoS Med</i> 2008; 5 (2):e32 | 5 |
| 8 | Newton P, et al. Fake artesunate in southeast Asia. <i>Lancet</i> 2001; 357 (9272):1948-50 | 5 |
| 9 | Laserson K, et al. Substandard tuberculosis drugs on the global market and their simple detection. <i>Int J Tuberc Lung Dis</i> 2001; 5 (5):448-54 | 5 |
| 10 | ReMeD. La Qualite´ des me´dicaments sur le marche´ pharmaceutique africain: e´tude analytique dans trois pays: Cameroun, Madagascar, Tchad. Action Programme on Essential Drugs. In: WHO, ed. Geneva, 1995. | 5 |
| 11 | Baratta F, et al. Diffusion of counterfeit drugs in developing countries and stability of galenics stored for months under different conditions of temperature and relative humidity. <i>Croat Med J</i> 2012; 53 (2):173-84 | 5 |
| 12 | Seear M, et al. The need for better data about counterfeit drugs in developing countries: a proposed standard research methodology tested in Chennai, India. <i>J Clin Pharm Ther</i> 2011; 36 (4):488-95 | 5 |
| 13 | Amin AA, et al. The quality of sulphadoxine-pyrimethamine and amodiaquine products in the Kenyan retail sector. <i>J Clin Pharm Ther</i> 2005; 30 (6):559-65 | 4 |
| 14 | Odufa O, et al. Pharmaceutical Equivalence of Some Commercial Samples of Artesunate and Amodiaquine Tablets Sold in Southwestern Nigeria. <i>Trop J Pharm Res</i> 2009; 8 (6):491-99 | 4 |
| 15 | Kyriacos S, et al. Quality of amoxicillin formulations in some Arab countries. <i>J Clin Pharm Ther</i> 2008; 33 (4):375-79 | 4 |
| 16 | Pribluda V, et al. Implementation of basic quality control tests for malaria medicines in Amazon Basin countries: results for the 2005-2010 period. <i>Malar J</i> 2012; 11 (1):202 | 4 |
| 17 | Obodozie OO, et al. A comparative study on the prevalence of substandard ampicillin/cloxacillin preparations in the | 3 |

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|----|--|---|
| | Nigerian market: Mid 1990's and present. <i>Journal of Phytomedicine and Therapeutics</i> 2006; 11 (2006):1-8 | |
| 18 | Prazuck T, et al. Quality Control of Antibiotics Before the Implementation of an STD Program in Northern Myanmar. <i>Sex Transm Dis</i> 2002; 29 (11):624-627. | 3 |
| 19 | Bate R, et al. Antimalarial drug quality in the most severely malarious parts of Africa - a six country study. <i>PLoS One</i> 2008; 3 :e2132 | 3 |
| 20 | Atemnkeng MA, et al. Quality evaluation of chloroquine, quinine, sulfadoxine–pyrimethamine and proguanil formulations sold on the market in East Congo DR. <i>J Clin Pharm Ther</i> 2007; 32 (2):123-132. | 3 |
| 21 | Obaid A. Quality of ceftriaxone in Pakistan: reality and resonance. <i>Pak J Pharm Sci</i> 2009; 22 (2):220-9. | 3 |
| 22 | Abdo-Rabbo A, et al. The quality of antimalarials available in Yemen. <i>Malar J</i> 2005; 4 (1):28. | 3 |
| 23 | Roy J. The menace of substandard drugs. <i>World Health Forum</i> 1994; 15 :406-407. | 2 |
| 24 | Bate R, et al. Pilot Study of Essential Drug Quality in Two Major Cities in India. <i>PLoS One</i> 2009; 4 (6):e6003. | 2 |
| 25 | Iwuagwu MA, et al. In vitro assessment of ampicillin capsules marketed in Nigeria. <i>International Journal of Pharmacy Practice</i> 1992; 1 (3):167-171. | 2 |
| 26 | Abdullah M, et al. Report: in vitro dissolution studies of different brands of sustained release diclofenac sodium matrix tablet available in Bangladesh. <i>Pak J Pharm Sci</i> 2008; 21 (1):70-77. | 1 |
| 27 | Zaheer M, et al. In vitro Analysis and Data Comparison of Market Brands of Ciprofloxacin, Ofloxacin and Levofloxacin. <i>Pak J Sci Ind Res</i> 2009; 52 (4):186-190. | 1 |
| 28 | Alfadl A, et al. quality of antimalarial drugs in sudan: results of post-marketing surveillance. <i>Sudanese Journal of Public Health</i> 2006; 1 (2):108-111. | 1 |
| 29 | Kibwage IO, et al. Drug quality control work in Daru: observations during 1983-1986. <i>East Afr Med J</i> 1992; 69 (10):577-80. | 1 |

Table 4: The prevalence of counterfeit/substandard medicines.

| Country [Reference] | Drugs (n=number of various products tested) | Setting | Formulation studied | Labeled Origin | Method of testing/location* | Stated problems | % (substandard or counterfeit) | Methodological strength scoring (0-12) |
|--|---|--------------------------------------|----------------------|--|---|---|---|--|
| The prevalence of counterfeit and substandard medicines in low-income countries in Asia and Africa. | | | | | | | | |
| Lao PDR (17) | Ampicillin, tetracycline, Chloroquine and aspirin (n=300) | Private outlets | Tablets and capsules | Laos, Thailand, France and unknown origin. | HPLC, colorimetric test , ultraviolet spectrophotometry, thin-layer chromatography and mass uniformity analysis / National Food and Drug Quality Control Centre | No active Ingredient, Inadequate/ excessive active ingredient and mass uniformity failure | 22% (Substandard / Counterfeit) | 10 |
| Tanzania (16) | Antimalarial drugs (sulfadoxine-pyrimethamine, sulfamethoxy-pyrazine-pyrimethamine, amodiaquine, quinine, artemisinin derivative (n=304) | Public and private outlets | Tablets | Local and imported | HPLC and dissolution test with US pharmacopeia standards/ Ifakara Health Research and Development Centre, Tanzania | Dissolution failure, Inadequate active ingredient. | 12.2% (Substandard / Counterfeit) | 9 |
| Cambodia (15) | Antimalarial drugs (Quinine, artesunate, mefloquine, chloroquine and tetracycline) (n=451) | Public ,private and informal outlets | Tablets | 16 countries | HPLC, disintegration test, thin-layer chromatography and packaging analysis/ National Laboratory for Drug Quality Control (NLDQC) in Cambodia | Failed in dissolution or inadequate active ingredient. ,no active ingredient, wrong active ingredient | 27% (50/451 substandard and 72/451 counterfeit) | 6 |
| Uganda (18) | Chloroquine (n=92) | Private and informal outlets | Tablets, injection | Not stated | HPLC/ Makerere University laboratory | Inadequate/ excessive active ingredient | 44.5 % (Substandard / Counterfeit) | 6 |

| The prevalence of counterfeit and substandard medicines in low-middle-income countries in Asia and Africa | | | | | | | | |
|---|---|--------------------------------------|--|--|---|--|---|---|
| Indonesia (20) | Amoxicillin, chloramphenicol, ciprofloxacin, cotrimoxazole, tetracycline. (n=104) | Public ,private and informal outlets | Tablets, capsules | Indonesia | HPLC/ Farmalyse BV laboratories (certified laboratory registered in the European Union as a pharmaceutical control laboratory for chemical physical analyses) | Inadequate active ingredient | 18% (Substandard / counterfeit) | 8 |
| Nigeria (19) | Artesunate, dihydroartemisinin, sulphadoxine-pyrimethamine, quinine and chloroquine (n=225) | Public ,private and informal outlets | Tablets | Not stated | HPLC and dissolution test, US pharmacopeia standards were used/ London School of Hygiene and Tropical Medicine laboratory | No active ingredient, wrong active ingredient, inadequate active ingredient. | 37% (Substandard / Counterfeit) | 7 |
| Nigeria(22) | Antimalarial drugs, antibacterials, antituberculosis, anthelmintics and antifungals (n = 581) | Private outlet | Tablets, capsules, suspension and injection. | 12 countries (Europe, Asia and Africa) | HPLC and dissolution test, British Pharmacopeia standards were used/ The Robert Gordon University School of Pharmacy laboratories | Inadequate/ excessive active ingredient, no active ingredient | 48% (Substandard/ Counterfeit) | 6 |
| Cameroon (21) | Antimalarial drugs (Antifolates, quinine, chloroquine) (n=284) | Informal outlets | Tablets, capsules | Not stated | Thin-layer chromatography and Colorimetric test/ Unité de Recherche Paludologie Afro-tropicale, Institut de Recherche pour le Développement | No active ingredient, inadequate active ingredient, wrong ingredient, unknown ingredient | 39.4% (Substandard/ Counterfeit) | 6 |
| The prevalence of counterfeit and substandard medicines in the mixed group | | | | | | | | |
| Myanmar, Cambodia, Vietnam, Lao PDR, Thailand. (28) | Artesunate and mefloquine (n=232) | Public ,private and informal outlets | Tablets | China | HPLC, colorimetric testing (fast red dye) and packaging analysis/ Not stated | Fake packaging, no active ingredient | 44% (4 /232 substandard and 99/232 counterfeit) | 7 |

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|---|---|--------------------------------------|--|---|--|---|----------------------------------|---|
| Cameroon, Ghana, Kenya, Nigeria, Tanzania (26) | Antimalarial drugs (sulphadoxine-pyrimethamine, sulfamethoxypyrazine-pyrimethamine, artemisinin-based combination) (n=267) | Public ,private and informal outlets | Tablets | Local and imported (India, USA, Bangladesh, China, Mauritius, Vietnam and the UK) | Compendial quality testing according to US pharmacopeia standards/ WHO collaborating laboratory in South Africa | Inadequate/ excessive active ingredient, no active ingredient, mass uniformity, impurity and dissolution test failure | 28.5% (Substandard/ Counterfeit) | 7 |
| Uganda, Madagascar, Senegal (27) | Antimalarial drugs (Artemisinin-based combination, sulphadoxine-pyrimethamine) (n=188) | Public ,private and informal outlets | Tablets | Not stated | Compendial quality testing according to US pharmacopeia standards/ National Medicine Control Laboratory and laboratories at USP Headquarters | Dissolution failure, Impurity, Failure in the assay of active ingredient., mass uniformity test failure | 32% (Substandard/ Counterfeit) | 7 |
| Gabon, Ghana, Kenya, Mali, Mozambique, Sudan, Zimbabwe (24) | Antimalarial drugs (chloroquine and sulphadoxine-pyrimethamine) (n = 278) | Public ,private and informal outlets | Tablets, syrup | Local and Imported | HPLC, drug-specific c Assays and dissolution Test/ WHO collaborating laboratory in South Africa | inadequate active ingredient | 23% Substandard/ Counterfeit | 6 |
| Myanmar (Burma) and Vietnam (23) | Amoxicillin,ampicillin, metronidazole, paracetamol, salbutamol, tetracycline, chloroquine , chloramphenicol rifampicin and diazepam co-trimoxazole and ranitidine (n=500) | Public ,private and informal outlets | Tablets and capsules | More than 20 countries (Asia, Canada, Europe, USA and Australia) | Compendial quality testing according to British pharmacopeia standards/ WHO collaborating laboratory in Thailand | Inadequate/ excessive active ingredient, wrong active ingredient | 11% (Substandard/ counterfeit) | 6 |
| Nigeria and Thailand (25) | Chloroquine, amoxicillin, ampiclox cotrimoxazole, tetracycline, (n = 96) | Private and informal outlets | Tablets, capsules, suspension and injection. | Not stated | HPLC / Not stated | Inadequate/ excessive active ingredient | 36.5% (Substandard/ counterfeit) | 6 |

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|--|---------------------------------------|----------------------------|-------------------------------|--------------|--|--|---------------------------------|---|
| Armenia, Azerbaijan, Belarus, Kazakhstan, Ukraine, and Uzbekistan (29) | anti-tuberculosis medicines (n = 291) | Public and private outlets | Tablets, capsules, injections | 12 countries | HPLC, dissolution and mass uniformity test, US pharmacopeia standards were used/ Four WHO collaborating laboratories in Austria, Germany, Belgium and France | Content, mass uniformity, dissolution and related substances tests failures. | 11.3% (Substandard/counterfeit) | 6 |
|--|---------------------------------------|----------------------------|-------------------------------|--------------|--|--|---------------------------------|---|

HPLC: High-performance liquid chromatography; **USP:** United state pharmacopeia; **location*:** Location where is the analysis carried out.