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PHARMACEUTICAL COMPOUNDS NORMALLY ADDED IN ANTI- INFLAMMATION JAMU

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ABSTRACT

Background: Under Indonesian law, it is forbidden to add drugs to herbal medicine. Herbal medicine is commonly called Jamu. It is suspected that the direct efficacy experienced by herbal drinkers is due to the added harmful chemicals. **Objective:** To investigate forbidden pharmaceutical compounds in antiinflammation herbs circulating in the market. **Methods:** Analysis of forbidden pharmaceutical ingredients into jamu was performed using the standard color reaction, thin layer chromatography (TLC), and densitometry. Color reactions included organoleptic, solubility test and the color reaction, TLC using a stationary phase of silica gel GF 254 and a mixture of chloroform: acetone (80:20) eluent, then measured with a densitometer. **Results:** It was found from 7 antiinflammation jamu known on the market, three of which were added with synthetic compounds containing additional paracetamol and one of them was added with dexamethasone. **Conclusion:** There was a synthetic drug that was added to the herbal medicine namely jamu sold in the community. The obtained results suggest that the authorities more intensively to monitor the manufacture and distribution of jamu and herbal medicines.

Keywords: Jamu, Herbal medicine, Paracetamol, Dexamethasone TLC, Densitometer,

INTRODUCTION

In Indonesia, jamu or herbal medicine has been known since long as a traditional medicine processing and utilization are carried down through the generations by prescription heritage, beliefs, culture, and customs of this nation [1-5]. The words jamu derived from ancient Javanese word "jampi" (magic potion) implies the involvement of a magic spell by master traditional healing and the healing of disease by using magic spells or mantra. Herbs or herbal medicine is a traditional medicine that is widely used for the treatment because it is generally cheaper than synthetic drugs and allegedly did not give unwanted side effects as is the case with synthetic drugs [6,7]. According to the Regulation of the Minister of Health [8] of traditional medicine should not contain synthetic chemicals, the result of isolation is efficacious as drugs or narcotics, which are allowed as additives are substances that are not efficacious as a medicine. Jamu must be registered with the BPOM, Indonesian Agency for Food and Drug Administrations with special registration number.

The monitoring results in the market, there are many herbs which included indications that these herbs have medicinal properties, for example, antiinflammation drug doses resemble synthetic drug dosing. This has become one of the considerations of this study to determine whether there was a synthetic drug that was added to the herbal medicine sold in the community. Furthermore, the number of new medicinal plants popping up with a

product that is said to be nutritious "drink immediately healed" led to allegations addition of synthetic drugs into it. This article attempts to answer it.

MATERIALS AND METHODS Material

Materials used in the form of *jamu* antiinflammation from seven different companies, herbal medicine coded as A, B, C, D, E, F and G.

Color reagents used

Solution of sulfuric acid (98%) 18 M, a solution of 1% FeCl reagents, reagent Fehling's solution I, Fehling II and Nessler reagent solution. The color change before and after the addition of reagents is observed in accordance with the reference of Autherhoff [9].

Organoleptic

Organoleptic inspection to herbal required by Indonesian Pharmacopoeia, *Materia Medika Indonesia* and *Indonesian Herbal Pharmacopoeia* [10-16]. These observations included the dosage form, color, taste, and smell of herbs. The color observed after herbal homogenized, Inspection done with less taste tasting substances ways tip of the tongue and the smell examination performed by rubbing the substance on the fingertips and smelt.

Extraction

The extraction process additives in herbal medicine refer to the Minister of Health [16]. Herbal powder was added to the flask, added extractors which can dissolve the synthetic drugs that may be added to herbal medicine. Shaken until homogeneous and filtered, the filtrate was used for identification.

Reference standards

As a comparison which was expected to be added as forbidden compounds were antalgin, aspirin, paracetamol, mefenamic acid, ibuprofen, prednisone, and dexamethasone.

Thin layer chromatography (TLC)-densitometer

Extracts of jamu from the market and reference standards dissolved in certain solvents spotted on TLC plates with silica gel GF254. Let dry and then eluted using the eluent according to the synthetic

RESULTS AND DISCUSSION

Organoleptic

The composition of each of the jamu used was as follows, jamu A (piperis nigri fructus 15%, massoia aromatic 15%, cinnamomi cortex 10%, zingiberis rhizome 25%, abri folium 20%, orthosiphonis folium 10%, and others 5%), jamu B (zingiberis cassumunar rhizome 10%, curcumae rhizome 30%, zingiberis rhizome 30%, kaempferiae rhizome 20%, and myristicae rhizome 10%), Jamu C (kaempferiae rhizome 20%, curcumae rhizome 20%, retrofracti fructus 10%, zingiberis aromatica rhizome 15%, zingiberis purpureae rhizome 10%, parkiae cement

drugs were examined, namely, the eluent I (chloroform ethanol=90:10, eluent II (chloroform:acetone=80:20, and the eluent III (96% methanol: Concentrated ammonia (100:1.5). Observations were made under light ultraviolet 254 nm [10,17]. The dried plates were then put into an TLC- scanner and analyzed by densitometer. Peak area of chromatogram sample compared to the peak area standard multiplied by the number of milligrams of substance standard was weighed; the result was expressed in milligrams [18]. 15%, and others 10%), jamu D (myristicae cement 10%, zingiberis rhizome 30%, kaempferiae rhizome 20%, curcumae rhizome 30 and others 10%), jamu E (myristicae cement 10%, zingiberis rhizome 30%, kaempferiae rhizome 20%, curcumae rhizome 30%, retrofracti fructus 10%), jamu F (zingiberis rhizome 20%, curcumae domesticae rhizoma 20%, retrofracti fructus 15%, and others 25%), jamu G (kaempferiae rhizome 20%, 20% curcumae rhizome, retrofracti fructus 15%, 5% zingiberis aromatica rhizome, rhizome purpureae zingiberis 15%, and cement parkiae 15%).

Organoleptic results are shown in Table 1.

Table 1: Organoleptic observations

Jamu	Shape	Color	Sense	Odor
A	Fine Powder	Chocolate	Bitter	Typical herbs
B	Powder	Yellowish-brown	Bitter mildly	Typical herbs
			spicy	
C	Fine Powder	Greenish brown	Bitter	Typical herbs
D	Powder	Chocolate	Bitter mildly	Typical herbs
			spicy	
E	Fine Powder	Yellowish-brown	Bitter	Typical herbs
F	Fine Powder	Yellowish-green	Bitter mildly	Typical herbs
			spicy	
G	Fine Powder	Chocolate	Bitter	Typical herbs

Solubility test

Solubility test was based on the nature of the polarity of each synthetic compound [18]. Solubility test results are shown in Table 2.

Color reaction

The presence of synthetic compounds can be identified by the use of reagents that can react with the compound being analyzed and will cause a reaction which we can observe that coloration, precipitation, and formation of gases or odors. Some of the color reaction is used for the identification of specific functional groups in the molecule, such as a color reaction with iron (III) chloride to produce

red to purple if there is an aromatic hydroxy acid, phenol, enol, pyrazolone, and phenothiazines; for example on aspirin, mefenamic acid, antalgine and paracetamol [9]. Color reaction could only be done on herbal medicine A, B and D because the yellow color of the herbs remains dissolved in the solvent water, acetone, ethanol and methanol. Color reaction at jamu A, B and D was undertaken to ensure the presence or absence of synthetic compounds (paracetamol). Other colors formed from the reaction were difficult to interpret because of the yellow color of the herbs were very disturbing observations.

Table 2: Results of solubility test

Substance	Water	Acetone	Ethanol	Methanol	Chloroform
Jamu A	+++	++	+++	+++	-
Jamu B	+++	++	+++	+++	-
Jamu C	+++	++	+++	+++	-
Jamu C	+++	++	+++	+++	-
Jamu D	+++	++	+++	+++	-
Jamu E	+++	++	+++	+++	-
Jamu F	+++	++	+++	+++	-
Jamu G	+++	++	+++	+++	-
Antalgine	+++	-	++	++	+
Mefenamic acid	-	+	+	+	++
Asesosal	+	++	+++	+++	++
Dexamethasone	-	+++	+++	++	-
Ibuprofen	-	+++	+++	+++	+++
Paracetamol	++	+++	+++	+++	+
Prednisone	+	++	++	++	+

+++ : Soluble, ++ : Slightly soluble, + : Poorly soluble, - : Hardly soluble

TLC-densitometer

TLC carried out on the vessel that had been saturated with mobile phase. The stationary phase used in the form of silica gel GF 254 containing indicator fluorescence (sulfide inorganic) that

would emit light when illuminated at a wavelength of 254 nm. Spotting process was done carefully to obtain round and compact spots. The procedure followed the general reference of TLC [19-21].



Fig. 1 shows the typical TLC plates used.

The TLC plates were then measured in densitometer as TLC scanner. Measurement based on the reduction of the signal between the light transmitted or reflected by the plate blank part with areas containing the sample. Compounds being analyzed will be able to fluoresced and measured [22-25]. Chromatogram peak area of substances measured in comparison to the standard substance peak and then multiplied by the number of milligrams of standard substances was weighed and expressed in milligram [26,27]. Measurements were taken at maximum wavelength of 252 nm. A standard curve was made for each plate on level calculation was calculated

as the methods spektrofotometry with linear equations. Table 3 shows the results of analysis of illicit substances in Jamu samples. In this research, the yield returns of 97.08% and 98.01% for paracetamol and dexamethasone, respectively.

The use of jamu is usually recommended 3 times a day with brewed with 250 ml of water each sachet, meaning if a pack of herbal antiinflammation weight of 7 g of the herbal medicine contains approximately 450 mg of paracetamol. This is likely to cause the user or customer to think that she was taking the herbs that are effective with not aware of any additional substances into it.

Table 3: The results of the analysis of forbidden substances in Antiinflammation Jamu

Sample	%Forbidden compounds	
	Paracetamol	Dexamethasone
Jamu A	7.5	
Jamu B	6.8	
Jamu C	-	
Jamu D	8.0	
Jamu E	-	
Jamu F	-	
Jamu G	-	8.5

Paracetamol is a painkiller (analgesic) which is actually safe if used according to rules. These drugs are often found in cold medicine component and headache. When mixed into herbs, such as herbs or herbal pegel rheumatic arthritis, it will increase the efficacy of herbal medicine. If only used once or twice is not harmful to health. However, the problem is people generally assume that herbal medicine is safe and they tend to consume every day. If used every day, then paracetamol will accumulate in the body. In large doses, paracetamol can damage the liver/liver causing liver disorders. In the body, will paracetamol is metabolized to produce substances called free radicals N-acetyl-p- benzoquinoneimine (NAPQI). Under normal circumstances, would NAPQI didetoksikasi rapidly by the enzyme glutathione from the heart. In excessive doses, the liver can no longer mendetoksikasinya, and oxygen free radicals can actually damage the liver [28]. Prohibition dexamethasone associated with the following. Dexamethasone is drugs known as corticosteroids who has strong anti-inflammatory effects. If used according to the rules, corticosteroid drugs have many uses, especially in inflammatory diseases, such as rheumatoid arthritis, asthma, allergies, and autoimmune

diseases such as lupus, nephrotic syndrome, and rheumatoid arthritis. However, on the other hand, side effects are quite extensive, among others, is to increase blood sugar levels (increasing the risk of diabetes), bone loss (osteoporosis), inhibits the growth of children, cause fat in certain body parts (round face/ moon face, shoulders like humped), causing red streaks in the abdomen (striae), lowered immune system so prone to infection, increasing the risk of hypertension due to hold the salt in the body, causing stomach upset (stomach bleeding), etc. The use of these drugs should not be arbitrary and needs the supervision of the health authorities. If used as directed, of these, side effects can be managed so as not to endanger. Any cessation must not be sudden because it can also be harmful to health. Why is that? Since during the use of corticosteroids from the outside, the production of these hormones naturally from the body stops. If the use of outside suddenly stopped, the body will lack this hormone are normal and will occur reactions are undesirable, such as blood sugar levels fall, the blood pressure drops dramatically from a sitting to standing (orthostatic hypotension), dehydration (lack fluid), weakness, and lethargy. Uncontrolled usage in the form of herbal

medicine can certainly increase health risks for patients. Instead of a healthy, even admitted to hospital due to the use of herbs mixed with chemical drugs [29].

Our finding is inline with other researchers except for the methods and type of jamu used. Fauzi [30], for example, identified paracetamol in herbal medicine using TLC and Hafni [31] found their paracetamol in herbal medicine for the treatment of gout. Identification paracetamol in herbal fat healthy for men and women by thin layer chromatographic reported through thesis in 2009 by Siswanti [32]. The presence of paracetamol in herbal stiff circulating in Purwokerto, Mid Java, transparently was reported by Firdaus and Utami [33]. The addition of paracetamol in herbal medicine for uric acid was also found in Samarinda (West Borneo) as to which were reported in 2014 by Nurhasnawati *et al.* [34]. in Surabaya, East Java, was found the addition of paracetamol and phenylbutazone in herbal medicine stiff reported through thesis by Tourisma in 2011 [35]. By 2015 the identification of the kortikosteroid (dexamethasone, fenilbutazon, and prednisone) content in herbal medicine muscle pains reported by Widyawati *et al.* [36]. Addition of methampyrone, prednisone, and dexametazon was reported in Serang Province [37]. The results of this study add to the data of other researchers and have proven that there are still many illegal substances were added to the circulating medicinal herbs which are not in accordance with Permenkes No. 246/Menkes/Per/V/1990 although BPOM announced that 54 traditional medicines containing

forbidden compound [38]. In constrast to our study, Rukoyah *et al.* claiming RP-HPLC may be used for simultaneous analysis of four markers, namely, scopoletine, andrographolide, quercetin, and luteolin in antihypertension jamu. The methods can be used to analyze antihypertension jamu formulation in the market [39]. On the other hand, Wisnuwardhani *et al.* [40] proposed simultaneous analysis of steroid and nonsteroid anti-inflammatory substances in Jamu pegal linu using TLC-spectrophotometry.

We predicted that the addition of paracetamol and dexamethasone were possible because the price of these compounds relatively cheaper and easy to find than other antiinflammation raw materials in Indonesia.

CONCLUSION

There was a synthetic drug that was added to the herbal medicine namely jamu sold in the community. Of 7 antiinflammation jamu examined, three were found to the addition of paracetamol, and the addition dexamethasone on one type of them. This report suggests that the authorities more intensively to monitor the manufacture and distribution of jamu and herbal medicines.

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