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Focus On the Use of Shea Butter as Excipient For Ointment

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ABSTRACT

Shea butter is a fat produced from the seeds of *Vitellaria paradoxa* (*Sapotaceae*). It can exist under different qualities which depend on several factors such as the region of origin, the method of production, treatments performed, etc. Its use as ointment excipient was the subject of several work that has raised issues that need further discussion. In this study, certain aspects relating to the adequacy between the main properties required for the ointment excipients and the characteristics of the different qualities of shea butter were analyzed. The data collected showed that for some properties, namely organoleptic characteristics such as color and odor, it may be necessary to perform discoloration or deodorization operations to ensure the comfort of the user. Similarly, it has sometimes been necessary to add peanut oil to the butter to reduce its consistency in order to facilitate its spreading. Concerning the release of the active ingredients, the qualities of the butter seem to have no influence because whatever the butter used, the results obtained were satisfactory. As regards inertia and long-term stability, the composition could have a great influence, but the refined shea butter obtained by chemical extraction seems more apt to act as an excipient. However, for such a product, it is still necessary to standardize industrial procedures for obtaining it, with a view to propose a standard shea butter, specifically for use as ointment excipient.

Keywords: refined shea butter, unrefined shea butter, excipient for topical forms, ointment

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INTRODUCTION

Shea butter is a fat produced from the seeds of *Vitallaria paradoxa* (Sapotaceae). It is widely used in various sectors such as cosmetology (1-5) but especially in food industry (6-8) which would consume about 90% of world production (9, 10).

One of its peculiarities is the great variability of its quality. Indeed, there are several types of shea which are distinguished by different physicochemical profiles. This variability depends on several factors such as the area of origin, method of production, treatment implemented or storage conditions (11-16).

Its use in the pharmaceutical field, although being quantitatively less important, nevertheless attracts much interest. This is justified, in large part, by its pharmacodynamic properties, in particular anti-inflammatory properties associated with its unsaponifiable fraction (17- 22). But apart from its therapeutic properties, shea butter has other potentialities that have interested researchers. Indeed, its excipient properties have been the subject of numerous works, some of which date back several decades (23-26). Subsequent studies have confirmed such properties (27, 28). It should be noted, however, that in these studies, some aspects have been little or not addressed. These concern in particular, questions relating to the adequacy between the properties required for the excipients for topical forms and the quality of the butter used; the present work aims to analyze these issues in order to provide elements to better guide the choice of shea butter to use as an excipient when formulating ointments.

Some reminders about the required properties of excipients for topical use

The excipients used in topical forms, especially for ointments are defined on the one hand by physicochemical characteristics described in detail in the pharmacopoeias and, on the other hand by technological characteristics to meet the requirements for the production and the conditions of use of such forms.

They must have certain qualities, the main ones of which are set out below (29-32):

- A consistency allowing an easy spreading on the skin.
- Organoleptic characteristics (odor and color) that do not bother the user
- Absence of irritation, good tolerance and weak allergenicity
- Miscibility to the aqueous or fatty secretions of the skin. Thus, it can influence, according to its nature, the speed of penetration of an active principle capable of crossing the cutaneous barrier,
- Sufficient stability to allow good preservation

- Inertia towards the active ingredient (the excipient must not inhibit or increase the activity of the active ingredient), the packaging material (the excipient must neither dissolve the packaging material nor be absorbed by it), the body (in principle, the excipient must have no own activity or at least a pharmacological action compatible with that of the active ingredient).
- The ability, if possible, to be washable with water and not stain the laundry
- The possibility of being sterilized in the case of ointments to be applied to wounds.

Methods of obtaining and variations of the quality of shea butter

Shea butter can be produced by several methods (traditional, artisanal, industrial); these may have many modalities in their implementation and can be followed or not by complementary operations such as refining

However, despite these numerous parameters which constitute factors of variation in their quality, the butters obtained can be classified into two types essentially: butter extracted with or without solvent and butter refined or not after extraction. According to these criteria, the composition of the butter may sometimes vary significantly. Table I shows the difference in composition which may exist between two butters which have been extracted with hexane but one was subsequently refined and the other not (33).

Table I: Characteristic of unrefined and refined shea butter (33)

Characteristics	Crude oil (not refined)	Refined oil
Saponification value (mg KOH/g)	389.89	162.61
pH	4.38	4.58
Free fatty acid (%)	4.28	1.68
Acid value (mg KOH/ g)	8.42	3.36
Specific gravity	0.86	0.89
Moisture content (%)	2.29	0.12
Peroxide value (meq/kg)	15	9.40
Refractive index	1.472	1.467

In another study (34), two unrefined butters, one of which was extracted with water and the other with hexane, were compared and the results also showed differences for most measured parameters (Table II)

Table II: Physico-chemical Properties of SBT and SBS (34) (SBT: Shea butter extracted using traditional method; SBS: Shea butter extracted using solvent (hexane) extraction method.

Properties	SBT	SBS
Relative density	0.908	0.851
Kinematic viscosity (mm ² s ⁻¹)	30.68	44.84
Melting point (°C)	33.0	40.5
Free fatty acid (%)	9.0	23.89
Peroxide value (mg/100 g)	29.5	44.01
Acid value (mgKOH/g oil)	21.85	48.63
Iodine value (gI ₂ /100 g oil)	21.43	13.19
Saponification (mgKOH/g oil)	167.4	202.9
pH value	6.09	5.02

The unrefined butter itself may have several qualities which are distinguished by different physicochemical properties.

Thus, the West African Regional Standard (35) defined three grades of unrefined shea butter (Table III)

Table III: Quality Characteristics and Grading System for Unrefined Shea Butter (35)

Parameters	Grade 1	Grade 2	Grade 3
	<i>Max – Min</i>	Max-Min	Max-Min
Moisture content (%)	0 - 0.05	> 0.05 - 0.2	> 0.2 - 2.0
Free fatty acid (%)	0 - 1.0	> 1.0 - 3.0	> 3.0 - 8.0
Peroxide value (meq/kg)	0 - 10.0	>10.0 - 15.0	> 0.2 - 2.0
Insoluble impurities (%)	0 - 0.09	> 0.09 - 0.2	> 0.2 - 2.0

Organisms such as the American Shea Butter Institute (ASBI) have proposed other classification criteria that define five qualities of butter: A (raw or unrefined, extracted using water), B (refined), C (highly refined and extracted with Solvents such as hexane), D (lowest uncontaminated grade), E (with contaminants) (36).

Critical analysis of the use of butter in the formulation of ointment

Whatever the standard of classification, the different types of butter defined therein are characterized by physicochemical properties that justify their use in specific areas. According to the Regional Technical Committee, grade 1 unrefined shea butter is intended for the cosmetics and / or pharmaceutical industries and for direct consumption, the second category (grade 2) is intended for the food industries (confectionery, chocolate factories , kitchen and margarine ...), while the third category (grade 3) is for soap and direct consumption after refining (35)

However, as regards its use as excipient for ointment, a well-defined quality does not seem to be specified. It is therefore legitimate to ask the following questions: do all types of butter possess the main properties required for topical excipients? If not, for which properties would one or the other type be more apt to play the role of excipient?

Color and odor

Color varies according to the region of origin, the quality of the nuts and the conditions of production (37); It can thus vary from pale yellow to yellow (38, 39, 40) which is considered as that of unrefined butter. The refinement loses this original color which becomes whitish (41, 42).

Raw shea butter has a characteristic odor that some find unpleasant. The latter, which is more pronounced for shea butter in West Africa compared with that from East Africa, can however be corrected by various deodorization and aromatization processes (43).

Consistency and release of active ingredients

In publications where shea butter was used as excipient, the main objectives of the authors were the possibility of preparing an ointment whose consistency allowed, on the one hand, good spreading and on the other hand the release of the active ingredients according to the desired profile. Raw unrefined shea butter has a relatively hard consistency which does not favor a good spreading power, whereas the latter is an essential quality required for ointments

Thus, to meet such a requirement, the use of shea butter as an excipient for ointment required the addition of a more fluid substance such as peanut oil at a concentration between 10% and 20% (23, 24, 26, 28) to have a suitable consistency capable of ensuring good spreading.

Its ability to release active ingredients has been studied in many works where shea butter has been compared with various other conventional ointment excipients. The results showed that the release profiles obtained with the shea butter were comparable to those observed with the reference excipients.

The release of Aureomycin from an ointment made with shea butter proved to be greater than when lanolin or petroleum jelly were used as excipients (28). Similar results were obtained when the release profiles of Metronidazole (44) and Sulfur (45) from shea butter based ointments and from Simple Ointment (BP) were compared. Similarly, salicylic acid was released faster from an ointment using shea butter as excipient than from the British Pharmacopoeia's reference ointment (26). In another study where shea butter was used as a base for an ointment containing Griseofulvin as active (46), the results showed a release of the latter with satisfactory profiles.

It should be noted that to meet these two criteria (good spreading and good release of the active ingredients with which it is associated), the type of butter used does not seem to be of particular

importance. Indeed, some have used butter purchased at the local market and then treated with different purification methods (28, 33). Sometimes there is no clarification of the treatment of butter before use (45, 47). In other cases, studies have focused on a comparison between purified butter and raw butter (46). But whatever the butter that was used, the technological qualities of the latter allowed the preparation of ointments which have been tested successfully.

Inertia

With respect to the active ingredient

A reaction with the active ingredient could result in degradation of the latter leading to its inactivation or toxicity for example (48-52). Publications where shea butter has been used as an excipient for various active ingredients do not mention such a phenomenon (26, 28, 44, 45).

With respect to the organism

This is one of the most important properties for an excipient. Nowadays, the activity of certain excipients towards the organism is increasingly mentioned, but in general reference is made to excipients with a known effect (53-55). For topical products, allergic or irritation reactions (54, 55) can sometimes occur but with shea butter-based ointments, such reactions have not been reported. On the contrary, with unrefined shea butter, there is rather a beneficial effect that is noted. Indeed, apart from its importance in the agro-food industry, unrefined shea butter owes its current popularity to the many cosmetic and therapeutic virtues attributed to its unsaponifiable fraction that is rich in active principles, in particular anti-inflammatory (17-22). Therefore, if unrefined shea butter is to be used, the activity of the active ingredient to which it is to be combined must be different from that of the principles contained in the unsaponifiable. If this is not the case, the evaluation of the active ingredient could be biased. For example, it would be difficult to distinguish between the activity of an anti-inflammatory active ingredient and that due to the unsaponifiable. Ointments prepared with the barks of *Khaya senegalensis* (Meliaceae), an anti-inflammatory plant, using shea butter as a carrier (56-58), could be cited to illustrate such a phenomenon. Indeed, the anti-inflammatory activity of such ointments can not be attributed only to the barks of the plant.

As regards refined shea butter, the treatments which led to its obtaining destroyed most of the active ingredients contained in its unsaponifiable fraction (59, 60); thus, it would better meet the criterion of inertia required vis-à-vis the organism.

Long-term stability

Various parameters such as moisture content, insoluble impurities, free fatty acids and peroxides can lead to fatty alteration reactions and thus, limit their shelf life. Among these reactions, the

oxidation, which is responsible for the rancidity phenomenon, represents one of their principal degradation pathways (61- 66).

The substrates of the oxidation reactions consist essentially of the free fatty acids. Unrefined shea butter being much richer in free fatty acids (see Table I), would therefore be more sensitive to such reactions.

It should also be noted that these oxidative phenomena are accompanied by a deterioration of the organoleptic characteristics such as the occurrence of an unpleasant odor (63). However, when stored in good conditions, these reactions can be delayed by, in part, the natural antioxidants found in the unsaponifiable fraction (7, 3). But this would only be possible if unrefined shea butter is obtained in the absence of a high temperature capable of destroying these antioxidants.

In order to improve its stability, it would therefore be necessary to associate it with antioxidants (24, 67).

As for refined shea butter, the operations allowing its obtaining lead to the elimination of a large part of the free fatty acids and thus constitute a factor of stability (3, 67, 68) favorable to better conservation over time. However, what is important is the maintenance of the stability of the ointment using the shea butter (refined or not). A dosage of the various components just after the preparation of ointments and after a defined time could allow an objective judgment.

In total, when comparing unrefined butter and refined butter, it can be seen that both can be used indifferently if one considers the spreading power and the ability to release the active ingredients. However, as regards the other properties required for excipients for topical use, such as the ability to ensure user comfort, inertia to the body and long-term stability, refined shea butter seems more apt to play the role of excipient. The availability of such a product, however, comes up against a difficulty often found in the world of excipients, namely the lack of industry that is specifically intended for them (69, 70). In general, the excipients used in the pharmaceutical formulation are also produced for other purposes. Shea butter most suitable for use as an ointment excipient is that produced by the food industry because it has the essential properties required for topical application (moisture content, free and unsaponifiables fatty acids are low, the color is stable and there is no odor) (3). However, the procedures used should be adapted to the requirements for a pharmaceutical use (71).

CONCLUSION

This analysis has shown that it is possible to use different types of shea butter as an excipient for ointment but in some cases it has been found necessary to modify certain characteristics of the

butter in order to make them conform to the properties required for such use. These modifications are justified by the fact that there is not on the market a standard shea butter especially intended for use as an excipient in the pharmaceutical formulation of ointment. Nevertheless, it may be noted that the characteristics of refined shea butter seem more suited to the properties required for the excipients for topical use. However, the procedures for preparing and distributing such a butter should be standardized in order to solve the problems associated with the choice of the butter to be used and, moreover, to ensure the reproducibility of the formulations in which it is to be incorporated.

REFERENCES

- 1- Alvarez AMR, Rodríguez MLG. Lipids in pharmaceutical and cosmetic preparations *Grasas y Aceites* 2000; 51 (1-2): 74-96.
- 2- Abbiw D K. Useful plants of Ghana, West Africa. Uses of wild and cultivated plants. Intermediate technology publication and the royal botanic gardens. London: Kew; 1990: 66-7.
- 3- The shea butter value chain production, transformation and marketing in West Africa. WATH Technical report, USAID West Africa 2004; N° 2.
- 4- Alander J. Shea Butter- a Multi Functional Ingredient for Food and Cosmetics. *Lipid Technol* 2004;16:202-205.
- 5- LE TRAITEMENT, LA VALORISATION ET LE COMMERCE DU KARITÉ EN AFRIQUE. Actes de l'atelier organisé par l' Organization des Nations Unies pour l'Alimentation et l'Agriculture, le Fonds Commun pour les Produits de Base et le Centre de Suivi Ecologique Centre de Suivi Ecologique. Dakar, Sénégal du 4 au 6 mars 2002: 232p.
- 6- Ogonnaya C, Adgidizi P. Evaluation of some Physico-chemical properties of Shea butter (*Butyrospermumparadoxum*) related to its value for food and industrial utilization. *International Journal of Post Harvest Technology and Innovation*. 2008; 1(3) : 320- 6
- 7- Bekure Z, Dolan M, Gordon Y, Tompson J. « Du local au mondial : le marché international du beurre de karité. » UNIFEM (Fonds de développement des Nations Unies pour la Femme), Dakar (Senegal) ; 1997 : 110 p.
- 8- Masters ET, Yidana JA, Lovett PN. Reinforcing Sound Management through Trade: Shea Tree Products in Africa. *Unasyva*2004; 210: 46-52.
- 9- USAID (2010). Nicole R. Investing in Shea in West Africa West Africa Trade Hub (WATH) Technical Report March 2010.

- 10- BEURRE DE KARITÉ : Comprendre la difference entre un beurre de karité naturel et un beurre de karité raffiné ; Guide Technique COSMEBIO, September 2014.
- 11- Di Vincenzo D, Maranz S, Serraiocco A, Vito R, Wiesman Z, Bianchi G. Regional variation in shea butter lipid and triterpene composition in four African countries. *J Agric Food Chem* 2005; 53(19):7473-9.
- 12- Maranz S, Weisman Z, Bisgaard J, Bianchi G. Germplasm resources of *Vitellaria paradoxa* based on variation in fat composition across the distribution range. *Agroforestry Systems* 2004; 60: 71-76.
- 13- Sonau H, Picard N, Lovett PN, Dembele M, Korbo A, Diarisso, Bouvet JM. Phenotypic variation of agromorphological traits of the shea tree, *Vitellaria paradoxa*. C.F. Gaertn., in Mali. *Genetic Resources and Crop Evolution* 2006; 53: 145-61.
- 14- Mbaiguinam M, Mbayhoudel K, Djekota C. Physical and Chemical Characteristics of Fruits, Pulps, Kernels and Butter of Shea *Butyrospermum parkii* (Sapotaceae) from Mandoul, Southern Chad. *Asian J Biochem* 2007; 2:101-10
- 15- Maranz S, Wiesman Z, Garti N. Phenolic Constituents of Shea (*Vitellaria paradoxa*) Kernels. *J Agric Food Chem* 2003b; 51: 6268-73.
- 16- Olaniyan AM, Oje K. Quality Characteristics of Shea Butter Recovered from Shea Kernel through Dry Extraction Process. *J Food Sci Technol* 2007; 44: 404-07.
- 17- J. Gassmüller. “Determination of Anti-Inflammatory Properties of Topical Formulations Containing Shea Butter Extract on Lesional Skin of Patients With Atopic Dermatitis;” BioSkin Institute for Dermatological Research, Hamburg, Germany; 2003.
- 18- Alander J, Andersson AC. The Shea Butter Family – the Complete Emollient Range for Skin Care Formulations. *Cosmetics and Toiletries Manufacture Worldwide* 2002: 28- 32.
- 19- Akihisa T, Kojima N, Kikuchi T, Yasukawa K, Tokuda H, Masters ET, Manosroi A, Manosroi J. Anti-Inflammatory and Chemopreventive Effects of Triterpene Cinnamates and Acetates from Shea Fat. *J Oleo Sci* 2010; 59 (6): 273- 80.
- 20- Nandini V, Chakrabarti R, Dass RH, Gautam HK. Anti-Inflammatory Effects of Shea Butter through Inhibition of Inos, Cox-2, and Cytokines via the Nf-Kb Pathway in Lps-Activated J774 Macrophage Cells. *Journal of Complementary and Integrative Medicine* 2012; 9(1):1-11.
- 21- Honfo FG, Akissoe N, Linnemann AR, Soumanou M, Van Boekel MA. Nutritional Composition of Shea Products and Chemical Properties of Shea Butter: A Review. *Critical Reviews in Food Science and Nutrition* 2014; 54: 673- 86.

- 22- Allal F, Piombo G, Kelly BA, Okullo JBL, Thiam M, Diallo OB, Nyarko G, Davrieux F, Lovett PN, Bouvet JM. Fatty acid and tocopherol patterns of variation within the natural range of the shea tree (*Vitellaria paradoxa*) Agro for Syst 2013; 87:1065–1082.
- 23- Kerharo J, Adam JG. La Pharmacopée sénégalaise traditionnelle. Plantes médicinales et toxiques. ed. Vigot Frères, Paris; 1974, 1012 p.
- 24- Mital HC, Dove FR. Shea butter as a base for ointments and creams. Drug and Cosmetic Industry 1973; 113(4):46-47 and 143-7.
- 25- Mital HC and Adotey J. Study of Shea Butter III: Comparative assessment of antioxidants and release of medicaments. Pharm Acta Helv. 1974; 49(1): 28-30.
- 26- Konning GH, Mital HC. Shea butter V. Effect of particle size on release of medicament from ointment. J Pharm Sci 1978; 67(2): 374- 6.
- 27- Oyedele AO. The skin tolerance of shea fat employed as excipient in topical preparations. Nig J Nat Prod Med 2002; 6: 26-9.
- 28- Thioune O, Khouma B, Diarra M, Diop AB and Lo I. The excipient properties of shea butter compared with vaseline and lanolin. J Pharm Belg 2003;58 (3) : 81-4.
- 29- Le Hir A, Chaumeil J-C, Brossard D, Charrueau C, Crauste-Manciet S. Pharmacie Galénique. Bonnes Pratiques de Fabrication des Médicaments. 10th ed., Issy-les-Moulineaux: Elsevier Masson; 2016 : 412.
- 30- Topical semi-solid dosage forms *The International Pharmacopoeia*. 9th ed., 2019.
- 31- Rajesh A, Sangeeta A, Deepak S, Prem CD, Nitin N. Topical ointment: an updated review. Journal of Drug Discovery and Therapeutics 2015; 3(25):47 - 51.
- 32- Usha S . Review on: an ointment. Ijppr. Human 2015 ; 4 (2) : 170-92.
- 33-** Sadiq MM, Musa U, Zinat A, Aris MI, Aliyu MA, Salihu Y. Extraction and characterization of Nigeria shea butter oil. Journal of Science, Technology, Mathematics and Education 2012 ; 8(2): 66-73.
- 34- Ajala EO, Aberuagba F, Olaniyan AM and Onifade KR. Optimization of solvent extraction of shea butter (*Vitellaria paradoxa*) using response surface methodology and its characterization. J Food Sci Technol 2016; 53: 730- 8.
- 35-** FAO/WHO COORDINATING COMMITTEE FOR AFRICA 22nd Session Nairobi, Kenya, 16 - 20 January 2017 PROPOSED DRAFT REGIONAL STANDARD FOR SHEA BUTTER. 2006: 6.
- 36- United States Agency for International Development. (2006) "Buying and Selling Shea Butter: A Marketing Manual for West Africa".

- 37-Chukwu O, Adgidzi PP. Evaluation of some physico-chemical properties of shea-butter (*Butyrospermum paradoxum*) related to its value for food and industrial utilization. Int J Postharvest TechnolInnov 2008;1: 320- 6.
- 38-Goreja WG. Shea Butter: The Nourishing Properties of Africa's Best-Kept Natural Beauty secret. Amazing Herbs Press, New York: 2004.
- 39-Moharram H, Ray J, Ozbas S, Juliani H, Simon J. Shea Butter: Chemistry, Quality, and new Market Potentials. In Wang M, Shengmin S, Hwang LS, Ho CT (ed).. Challenges in Chemistry and Biology of Herbal Research. Washington DC (USA): American Chemical Society, 2006; 326-40.
- 40- Sanogo R. Beurre de Karite en Dermopharmacie . AFEPHAR 4^{ième} Journée Scientifique Jeudi 02 Juin 2016.
- 41-Hamilton RJ, Rossell JB. Analysis of Oils and Fats. 1st ed., London, New York. Elsevier Applied Science: 1986.
- 42- Rossi M, Gianazza M, Alamprese C, Stanga F. The Effect of Bleaching and Physical Refining on Color and Minor Components of Palm Oil. J Amer Oil Chem Soc 2001; 78: 1051-5.
- 43- Ruiz Mendez MV, Huesa LJ, Pereda J, Dobarganes MC. Eusayos para la refinacion fisica de la menteca de karite. Grasas y aceites 1991, 42(2): 121- 6.
- 44- Oyedele AO. Delivery of Metronidazole From Purified Nigerian Shea butter In comparison to Standard And Modified Ointment Bases. Ife Journal of Science 2012; 14 (2): 253-7.
- 45- Mbang NF-O, Tolulope OA, Ayodele M. The Evaluation of Shea Butter from *butyrospermum parkii* as a Vehicle in Sulphur Ointment formulations. West African Journal of Pharmacy 2013; 24 (2): 58-65.
- 46- Ouoba B, Decroix O, Zuber M, Chaumeil JC. Feasibility Study of Dermatological Formulations based on Shea butter for the Tropical Countries of Africa. Drug Dev Ind Pharm 1994 ; 20(13) : 2091-2107
- 47- Dembélé DL. Formulation de pommade antalgique et anti-inflammatoire à base de *Securidaca longepedunculata* Fresen. Thèse de Pharmacie 2010-2011. Université de Bamako (Mali).
- 48- Sonali SB, Sandip BB, Amrita NB. Interactions and incompatibilities of pharmaceutical excipients with active pharmaceutical ingredients: a comprehensive review. J Excipients and Food Chem 2010; 1 (3): 3-26.

- 49- Nishath F, Tirunagari M, Husna KQ, Nandagopal A and Jangala VR. Drug-excipient interaction and its importance in dosage form development. *Journal of Applied Pharmaceutical Science* 2011; 01(06): 66-71.
- 50- Shilpa PC, Pradeep SP. Pharmaceutical Excipients: A review. *International Journal of Advances in Pharmacy, Biology and Chemistry* 2012;1(1): 21-34.
- 51- Kishore KH, Swapan R, VeerappanS. Drug-Excipient Interactions: Case Studies and Overview of Drug Degradation Pathways. *Am. J. Anal. Chem* 2016; 7: 107- 40.
- 52- Crowley P, Martini LG. Drug/excipient interactions. *Pharmaceutical Techn* 2001; 13: 26- 34.
- 53- Gérard J-P Placebo. Une question de principe. Mémoire D.I.U. Méthodologie de la Recherche Clinique et Epidémiologique. Univ. Antilles-Guyane Fac. Méd. 2002.
- 54- Fusier I., Tollier C-Y, Husson MC. Excipient à « effet notoire ». *Journal de pharmacie clinique* 2004 ; 23 : 149-56.
- 55- Liste des excipients à effet notoire. Mise à jour des libellés selon le Guideline Européen 2003. Afssaps- Deuxième révision du 03 Mars 2009.
- 56- Oumar Thioune, Jean Louis Pousset, Issa Lo. Mise au point d'une pommade à base de *Khaya senegalensis* : étude préliminaire et tests in vivo. XV^{èmes} Journées Médicales et Pharmaceutiques de Dakar, 17-22 Février 1997.
- 57- Thioune O, Ahodikpe D, Dieng M, Diop AB, Ngom S, Lô I. Pommade anti-inflammatoire à base de karité et d'extrait hydro-alcoolique d'écorces de *Khaya senegalensis* (Caïlcédrat). *Dakar Médical* 2000 ; 45 (2) : 113- 6.
- 58- Thioune O, Diouf A, Faye D, Lô I. Contribution a l'évaluation d'une pommade anti-inflammatoire à base de caïlcédrat (*Khaya senegalensis*). IV^{ème} Journée des Départements de la Section Pharmacie - Décembre 2007
- 59- Tasan M, Demirci M. Total and Individual Tocopherol Contents of Sunflower Oil at Different Steps of Refining. *Eur Food Res Technol* 2005 ; 220: 251- 4.
- 60- Van HV, Depaemelaere G, Villa AJ, Santiwattana P, Verhé R, De Greyt W. Influence of Chemical Refining on the Major and Minor Components of Rice Bran Oil. *J Amer Oil Chem Soc* 2006; 83: 315-21.
- 61- Frankel E. N. Lipid oxidation. *Prog. Lipid Res* 1980; 19: 1-22.
- 62- James C. Co-valorisation d'effluents grassex et résidus lignocellulosiques : déshydratation mécanique et compostage. Thèse Doctorat d'Université Limoges (France) n° 9 2007.
- 63- Réactions d'altération chimique des aliments. From: <http://www.azaquar.com/>. (accessed November 27, 2019 at 1pm).

- 64- Judde A. Prévention de l'oxydation des acides gras dans un produit cosmétique : mécanismes, conséquences, moyens de mesure, quels antioxydants pour quelles applications *OCL* 2004 ; 11 (6) : 414-8.
- 65- Jeantet R, Croguennec T, Schuck P, Brule G. Science des aliments- Biochimie-Microbiologie- Procédés-Produits. Vol.1. Stabilization biologique et physico-chimique. Tec&Doc (ed), Paris : Lavoisier, 2006.
- 66- Honfo F, Hell K, Akissoé N, Coulibaly O, Fandohan P, Hounhouigan J. Effect of storage conditions on microbiological and physicochemical quality of shea butter. *J Food Sci Technol* 2011; 48(3): 274- 9.
- 67- Seung NH, Juliani HR, Simon JE. Effects of Selected Synthetic and Natural Antioxidants on the Oxidative Stability of Shea Butter (*Vitellaria paradoxa* sub sp. *paradoxa*). *Journal of Medicinally Active Plants* 2012;1(2):69-75
- 68- Sub-saharan Africa : factors affecting trade patterns of selected industries, Second Annual Report, Inv. 332-477 Ed. Diane Publishing, April 2008.
- 69- Aurélie Dureuil. Excipients : un marché difficile à réglementer – Industrie Pharma N° 52 Décembre 2000.
- 70- Matières premières pharmaceutiques, Mondialisation et Santé publique ». Académie Nationale Française. Compte rendu Séance thématique du Mercredi 20 Avril 2011.
- 71- Qualification of Excipients for Use In Pharmaceuticals. The International Pharmaceutical Excipients Council 66 p.

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