

# DEVELOPMENT OF FLURBIPROFEN MICROEMULGEL

Sarah BOUAMEUR<sup>1\*</sup>, Fatma SAMEUR<sup>2</sup>, Rachida BENAOUAD<sup>2</sup>, Soumia CHIRANI<sup>2,3</sup>, Nassima MOUSSAOUI<sup>1</sup>

1 Laboratory of galenic pharmacy, Department of pharmacy. University of Oran 1, Algeria.

2 Department of pharmacy. University Djillali Liabes of Sidi Bel Abbès, Algeria.

3 Laboratory of Physic Macromolecular and Organic Chemistry. University Djillali Liabes of Sidi Bel Abbès, Algeria.

## Abstract

Topic application of Non Steroidal Anti Inflammatory Drugs (NSAIDs) presents the big advantage to avoid side effects caused by their oral administration. Among the topic forms, Microemulgel is a new topical dosage form thermodynamically stable and very interesting to enhance cutaneous permeability. Our objective was to formulate and characterize a microemulgel containing 5% of flurbiprofen.

**Methods** : Flurbiprofen microemulsion 5% was prepared by titration method with three excipients, olive oil, Tween 80 and Span 80. The adequate microemulsion proportions were determined in the pseudo-ternary diagram. Flurbiprofen was dissolved in the preparation by magnetic agitation. To obtain Flurbiprofen microemulgel, 1% of Carbopol 934 was added to the previous microemulsion. Microemulgel was characterized by transmittance, particle size, refractive index, conductivity, viscosity and stability was evaluated by freeze-thaw cycle test.

**Results** : Flurbiprofen Microemulgel 5% prepared was transparent (100% of transmittance), stable with low viscosity and pH. Globule size was characterized by  $104.4 \pm 66.13$  nm of diameter and refractive index was  $1.474 \pm 10^{-5}$ .

**Conclusion** : Microemulgel containing 5% flurbiprofen presents very interesting characteristics for topical administration. This innovation can provide more efficiency to treat rheumatism and inflammatory diseases.

**Key Words** : antiinflammatory, carbopol, formulation, microemulsion, topic application

## 1- Introduction

Rheumatic Inflammatory Diseases (RID) are very painful pathologies. They affect many people in the world and evolve to chronic forms with high morbidity. In Algeria, 50% of RID are responsible of fonctionnal difficulties [1]. The treatment is essentially based on Non Steroidal AntiInflammatory Drugs (NSAID) which lead to severe gastrointestinal side effects after oral administration. NSAID Topical forms represent a good alternative in this case, however their cutaneous absorption is very limited and articular zone cannot be reached [2,3].

To overcome these difficulties and to enhance the cutaneous permeability of NSAID, new galenic formulations are developed. Among innovative topic forms, microemulsion (ME) provides very interesting proprieties to increase cutaneous absorption. They consist of dispersed system with very narrow tenuity (10- 100 nm), characterized by thermodynamic and cinetic stability [4,5]. Their combinaison with an hydrogel lead to a microemulgel (MEG) form which provides more efficiency and stability to the ME dispersion [6].

In this work, we developed and characterized a MEG of flurbiprofen (FLU) 5% intended to anti rheumatic pathologies.

## 2- Material and method

Flurbiprofen Microemulgel (FLU-MEG) was developed according to theses different steps :

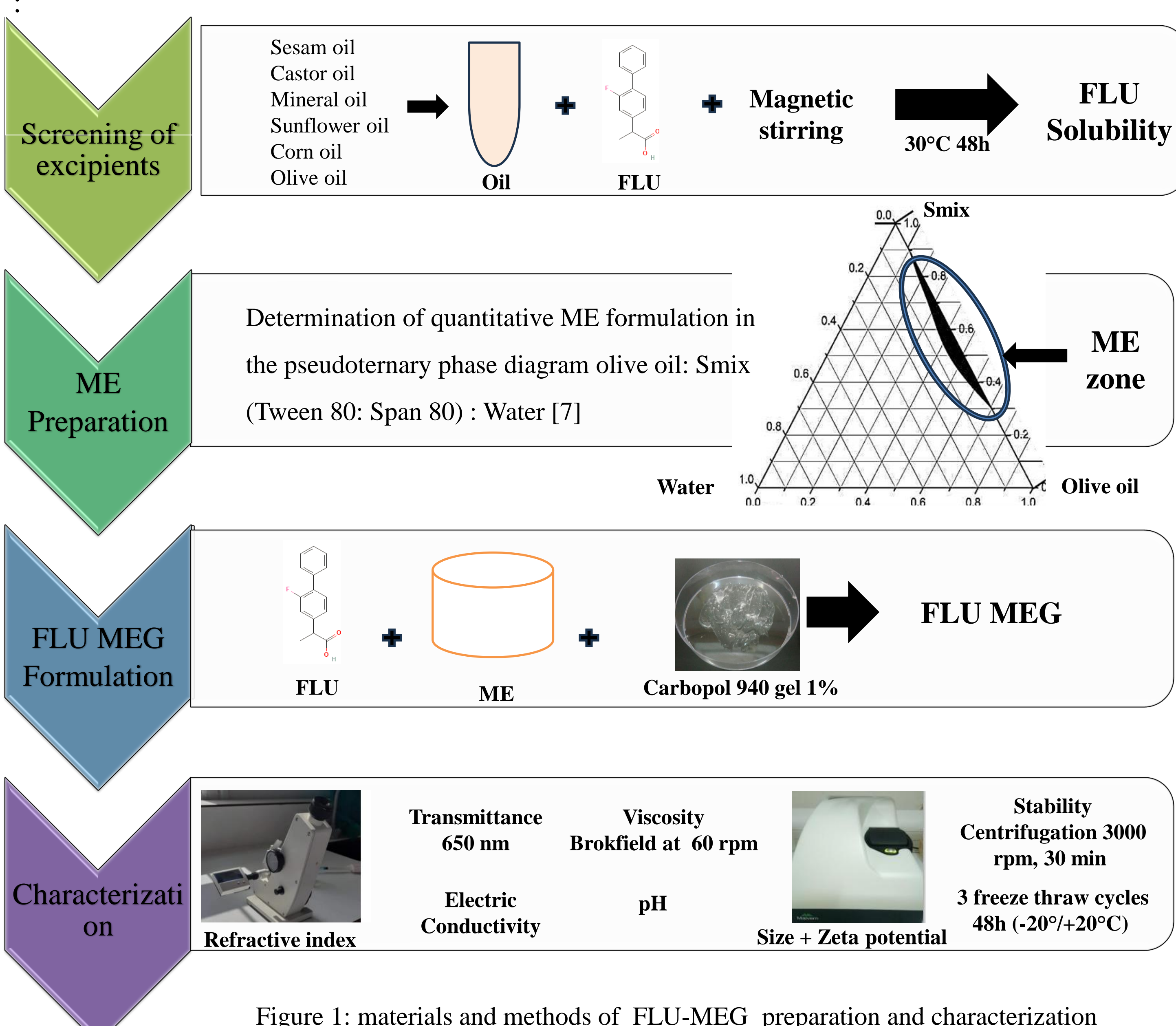


Figure 1: materials and methods of FLU-MEG preparation and characterization

## 3- Results & discussion

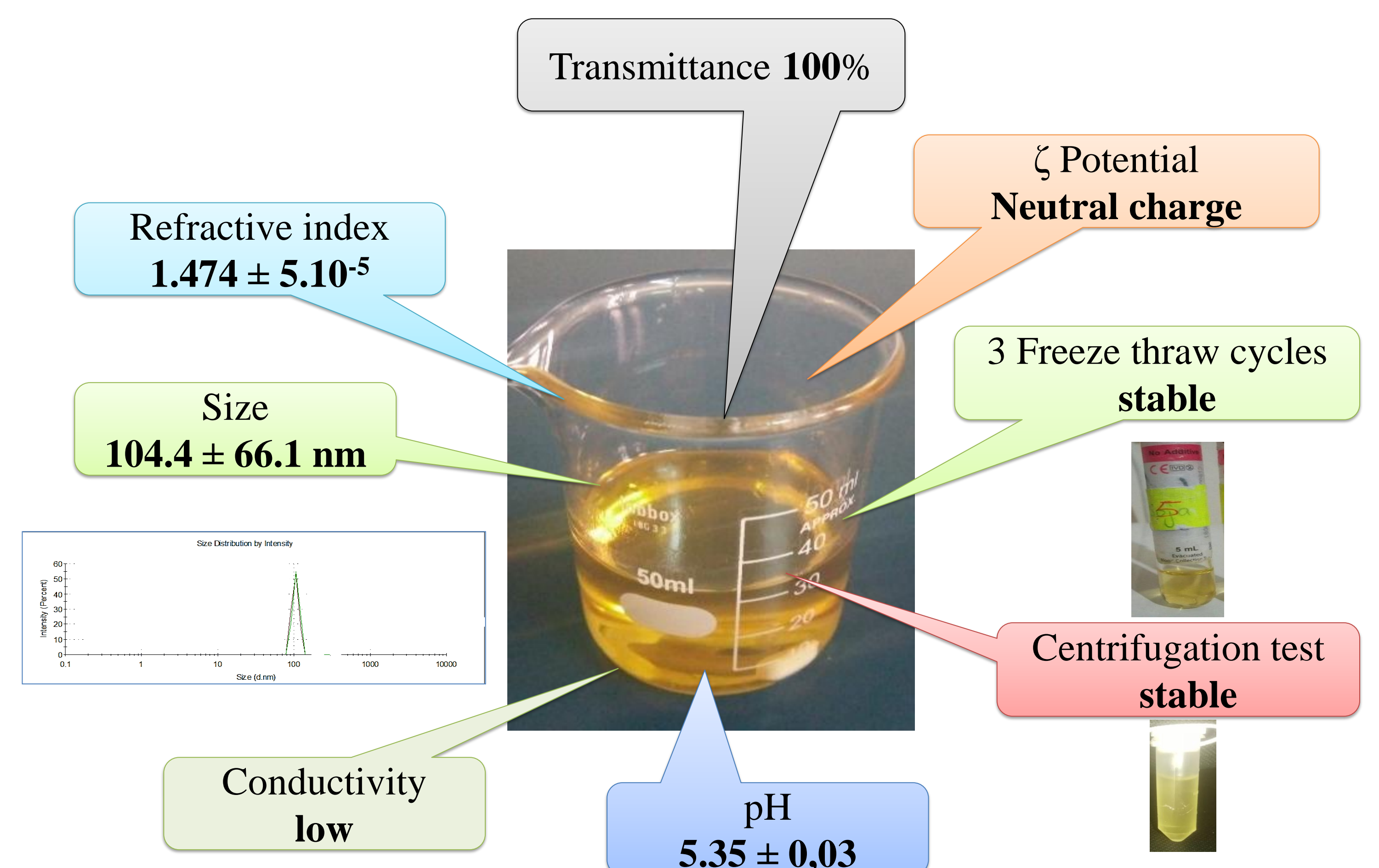


Figure 2 : results of ME and MEG of FLU 5%

Table 1 : compositions and some characteristics of FLU MEG in litterature

	FLU-MEG	[8]	[9]	[10]
Viscosity (cp)	350	108100	400	/
Composition	FLU 5% Olive Oil (56%) Tween 80 : Span 80 (1:1), Carbopol 940 NaOH Water	FLU 3% Arachis Oil (40%) Carbopol 934 Triethanolamine Water	FLU 5% Acid oleic (5%) Tween 20 : ethanol (2 :1) Carbopol 934 (46%) Triethanolamine Water (44%)	FLU 1% Clove oil (8.33%) Tween 80 : transcitol (33.33%) Carbopol 934 (1%) Triethanolamine Water

## 4. Conclusion

FLU-MEG 5% is very interesting form for anti inflammatory and anti rheumatic treatment. Formulation of stable and efficient MEG can be hard and laborious and needs complementary *invitro/ invivo* investigations to achieve FLU-MEG characterization.

## 5. References

- [1]. L'arthrite et les maladies rhumatismales inflammatoires : journée mondiale de l'arthrite 2012. [Internet]. 2012 [cité 8 mai 2023]. Disponible sur : <https://www.santetropicale.com/santemag/actus.asp?id=14986>
- [2]. Paudel KS, Milewski M, Swadley CL, Brogden NK, Ghosh P, Stinchcomb AL. Challenges and opportunities in dermal/transdermal delivery. Ther Deliv. 2010 Jul;1(1):109-31. doi: 10.4155/tde.10.16. PMID: 21132122; PMCID: PMC2995530.
- [3]. Barkin RL. Topical Nonsteroidal Anti-Inflammatory Drugs: The Importance of Drug, Delivery, and Therapeutic Outcome. Am J Ther. 2015 Sep-Oct;22(5):388-407. doi: 10.1097/MJT.0b013e3182459abd. PMID: 22367354
- [4]. Lawrence MJ, Rees GD. Microemulsion-based media as novel drug delivery systems. Adv Drug Deliv Rev. 6 déc 2000;45(1):89-121.
- [5]. Ghosh P, Murthy R. Microemulsions: A Potential Drug Delivery System. Curr Drug Deliv. 28 mars 2006;3(2):167-80.
- [6]. Ashara KC, Pann JS, Soniwala MM, Chavda JR, Mendapara VP, Mori NM. Microemulgel: an overwhelming approach to improve therapeutic action of drug moiety. Saudi Pharmaceutical Journal. 1 juill 2016;24(4):452-7.
- [7]. Yew HC, Misran M Bin. Nonionic Mixed Surfactant Stabilized Water-in-Oil Microemulsions for Active Ingredient in Vitro Sustained Release. J Surfactants Deterg. 1 janv 2016;19(1):49-56
- [8]. Ambade KW, Jadhav SL, Gambhire MN, Kurmi SD, Kadam VJ, Jadhav KR. Formulation and Evaluation of Flurbiprofen Microemulsion. Vol. 5, Current Drug Delivery. 2008.
- [9]. Naeem M, Rahman NU, Tavares GD, Barbosa SF, Chacra NB, Löbenberg R, et al. Physicochemical, in vitro and in vivo evaluation of flurbiprofen microemulsion. An Acad Bras Cienc. 1 janv 2015;87(3):1823-31.
- [10]. Battu H, Madhavi N, Reddy KN. Design, Development and in vitro assessment of Flurbiprofen Microemulsion Gel for Transdermal Drug Delivery. J Pharm Res Int. 2 août 2022;67-76.