

## Supplementary Material

# Optimization of Potential Nanoemulgels for Boosting Transdermal Glimepiride Delivery and Upgrading Its Anti-Diabetic Activity

Marwa H. Abdallah <sup>1,2,\*</sup>, Amr S. Abu Lila <sup>1,2,\*</sup>, Hanan M. El-Nahas <sup>2</sup> and Tarek M. Ibrahim <sup>2</sup>

<sup>1</sup> Department of Pharmaceutics, College of Pharmacy, University of Ha'il, Hail 81442, Saudi Arabia

<sup>2</sup> Department of Pharmaceutics, Faculty of Pharmacy, Zagazig University, Zagazig 44519, Egypt; hananelnahas@gmail.com (H.M.E.-N.); tarekmetwally333@gmail.com (T.M.I.)

\* Correspondence: mh.abdallah@uoh.edu.sa (M.H.A.); a.abulila@uoh.edu.sa (A.S.A.L.)

**Table S1.** Solubility of glimepiride in various nanoemulsion components.

Oils	Solubility (mg/g)
Peppermint oil	3.67 ± 0.13
Bergamot oil	2.75 ± 0.91
Mandarin oil	2.22 ± 0.19
Tea tree oil	2.11 ± 0.24
Eucalyptus oil	2.02 ± 0.17
Fennel oil	1.92 ± 0.26
Lavandar oil	1.40 ± 0.11
Rosemary oil	0.82 ± 0.10
Orange oil	0.50 ± 0.24
Lemon oil	0.43 ± 0.09
Surfactants	Solubility (mg/g)
Tween 80	2.94 ± 0.09
Tween 20	1.66 ± 0.21
Span 20	0.92 ± 0.19
Span 85	0.50 ± 0.13
Labrafil isostearique	0.61 ± 0.08
Labrafil 1944 CS	0.12 ± 0.03
Co-surfactant	Solubility (mg/g)
Transcutol P	2.54 ± 0.20
PEG 400	1.15 ± 0.14
PG	0.84 ± 0.29
Ethanol	0.81 ± 0.05
Lauroglycol FCC	0.32 ± 0.10
Glycerin	0.11 ± 0.05

**Table S2.** Miscibility of various surfactants and co-surfactants in oil phase.

Gel base type	Gel:NE ratio	Gel strength (sec)
Carbopol 940 (1%)	1.5:1	14.01 ± 0.23
	2.5:1	28.13 ± 0.84
Carbopol 940 (1.5%)	1.25:1	11.38 ± 0.88
	1.5:1	26.30 ± 1.02
Carbopol 940 (2%)	0.5:1	17.47 ± 0.10
	1:1	26.25 ± 0.91

	2:1	$8.54 \pm 1.12$
Carbopol 934 (1%)	3:1	$17.95 \pm 0.41$
	3.5:1	$25.26 \pm 0.86$
	1:1	$10.28 \pm 0.35$
Carbopol 934 (1.5%)	2:1	$18.84 \pm 1.40$
	3:1	$31.35 \pm 0.74$
Carbopol 934 (2%)	1:1	$15.82 \pm 1.21$
	1.25:1	$29.30 \pm 0.30$
	3:1	$12.89 \pm 0.91$
Na-CMC (5%)	3.25:1	$21.34 \pm 0.75$
	3.5:1	$31.85 \pm 0.29$
Sodium alginate (5%)	1:1	$13.39 \pm 0.35$
	1.25:1	$31.24 \pm 0.42$
	2:1	$10.02 \pm 0.96$
HPMC (5%)	3:1	$19.29 \pm 1.11$
	5:1	$27.35 \pm 0.82$

**Table S3.** Preliminary study for measuring gel strength values of different GM-loaded nanoemulgel formulations using different gel bases.

Oil/Surfactant	Number of additions (5 $\mu$ L)
Peppermint oil + (tween 80/water)	7 additions of oil
Peppermint oil + (tween 20/water)	6 additions of oil
Bergamot oil + (tween 80/water)	55 additions of oil
Bergamot oil + (tween 20/water)	50 additions of oil