



SUPPLEMENTARY MATERIAL TO
**Deep eutectic solvents formed by pharmaceutical ingredients
and their potential influences on solid preparations**

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Table S-I. VTF equation parameters of viscosity for the DESs.

DESs	A	B	T ₀	R ²
DES 1	3029.339	0	0.003595	1.0000
DES 2	3181.367	0	0.003595	0.9995
DES 3	2651.201	0	0.003595	0.9999
DES 4	3215.718	0	0.003595	0.9996

Table S-II. Excess molar volume of the menthol-citric acid DESs (5–35 °C).

Temperature	Excess molar volume			
	DES 1	DES 2	DES 3	DES 4
5 °C	-334.788	-345.165	-344.859	-349.867
15 °C	-335.122	-345.343	-350.652	-350.007
25 °C	-339.043	-346.275	-353.392	-352.382
35 °C	-354.165	-347.174	-355.207	-359.605

Table S-3. Excess molar volume of the phenyl salicylate-benzoic acid DESs (35–55 °C).

Temperature	Excess molar volume	
	DES 12:1	DES 9:1
35 °C	-283.749	-294.745
40 °C	-284.868	-295.417
45 °C	-285.253	-296.970
50 °C	-285.567	-297.425
55 °C	-286.202	-300.302

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Table S-IV. Solubilities of the DESs in different solvents at room temperature*

DESs	Solubility of DESs					
	Water	Glycerin	Ethanol	Methanol	Acetone	
menthol-citric acid	DES 1	PS	I	ES	ES	I
	DES 2	I	I	ES	ES	I
	DES 3	I	I	ES	ES	I
	DES 4	I	I	ES	ES	I
phenyl salicylate-benzoic acid	DES 12:1	I	I	ES	ES	I
	DES 9:1	I	I	ES	ES	I

*S: ES: easily soluble, I: insoluble and PS: poorly soluble

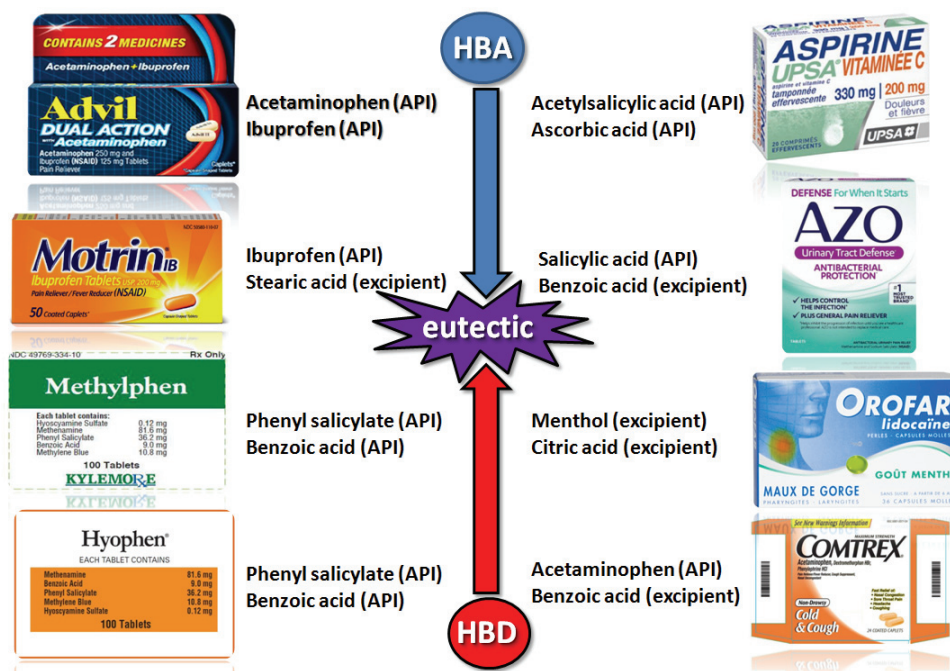


Fig. S-1. Common drug ingredients forming deep eutectic liquids.

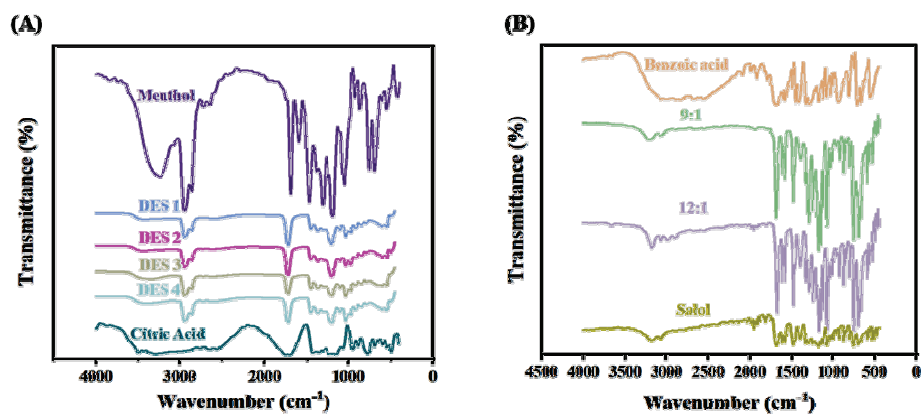


Fig. S-2. FTIR spectra of DESs composed of menthol and citric acid in various molar ratios (A); phenyl salicylate-benzoic acid DESs (B) and related pure components.

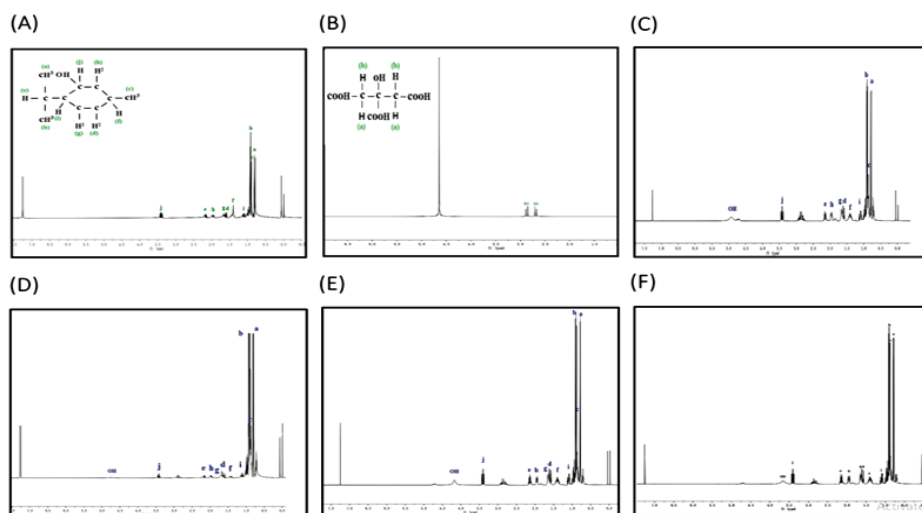


Fig. S-3. ¹H-NMR spectra (400 MHz, CDCl₃ for citric acid, DES 1-4 and D₂O for menthol) of (A) menthol, (B) citric Acid, (C) DES 1, (D) DES 2, (E) DES 3, and (F) DES 4.

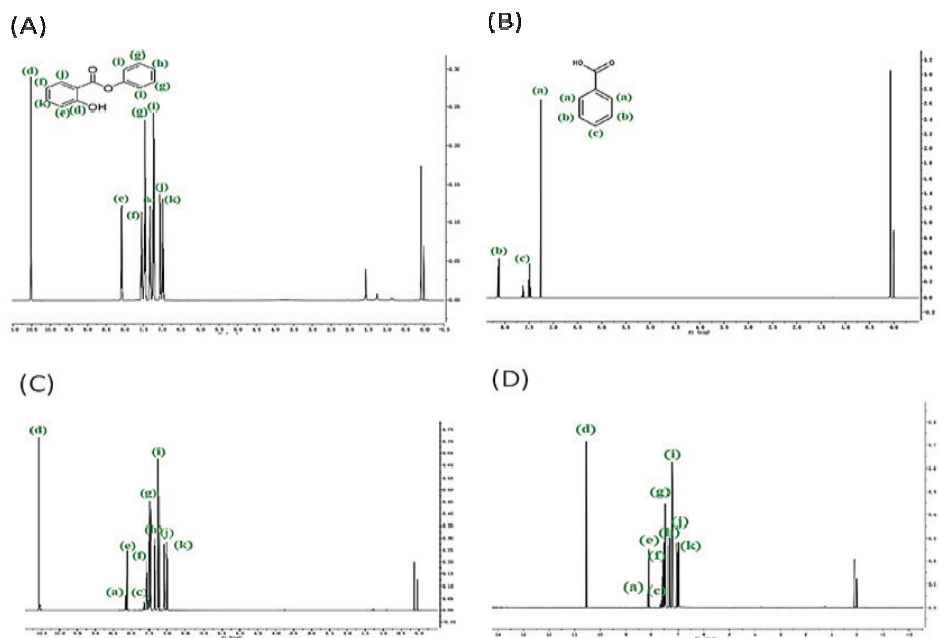


Fig. S-4. $^1\text{H-NMR}$ spectra (400 MHz, CDCl_3) of (A) phenyl salicylate, (B) benzoic acid, (C) their binary mixtures of 12:1 molar ratio, (D) binary mixtures of 9:1 molar ratio.

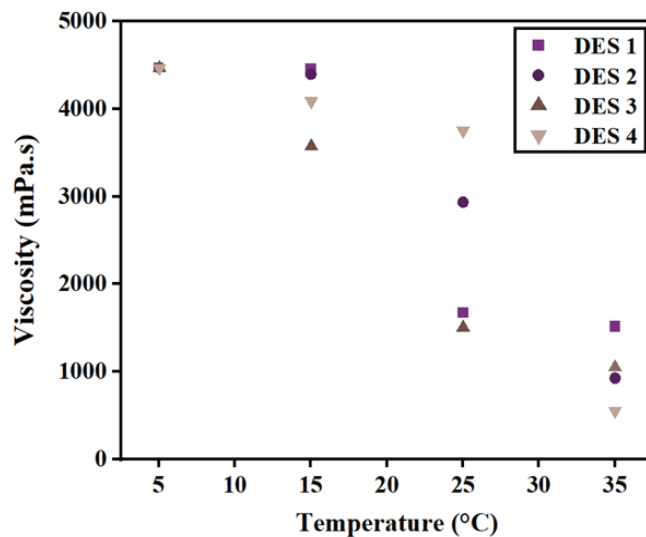


Fig. S-5. Plot of the predicted viscosity values of menthol-citric acid DESs.

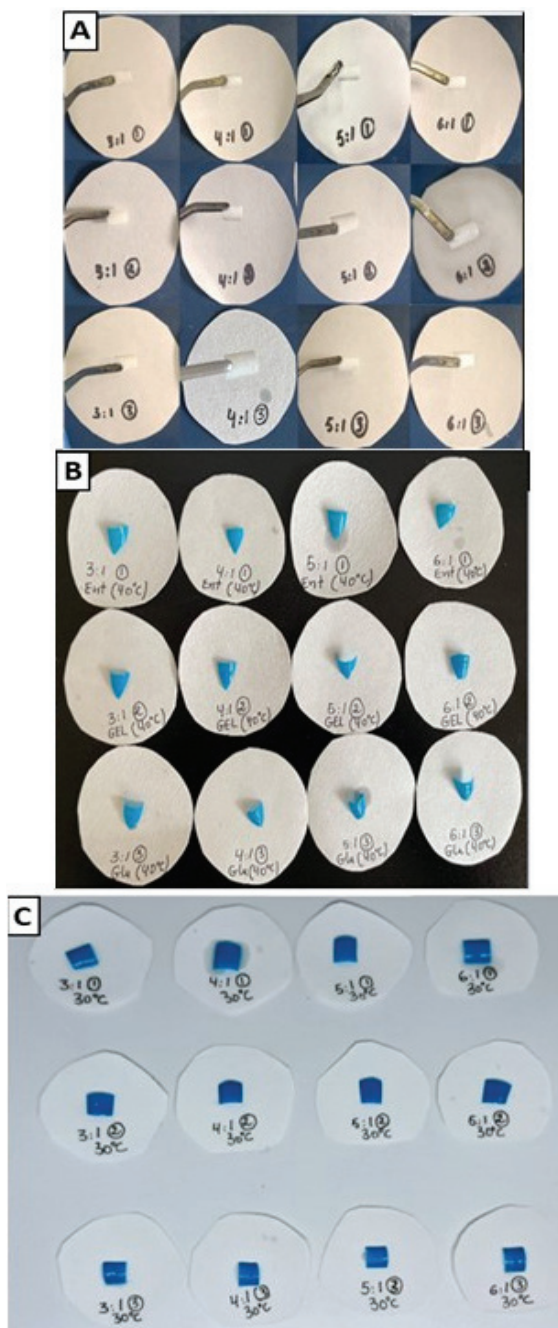


Fig.S-6. Photographs of the shells of enteric (1), gelatin (2) and glutinous (3) capsules taken after (A) 24 hours dissolved at 25°C; (B) 24 hours dissolved at 30°C; and (C) 24 hours dissolved at 40°C.

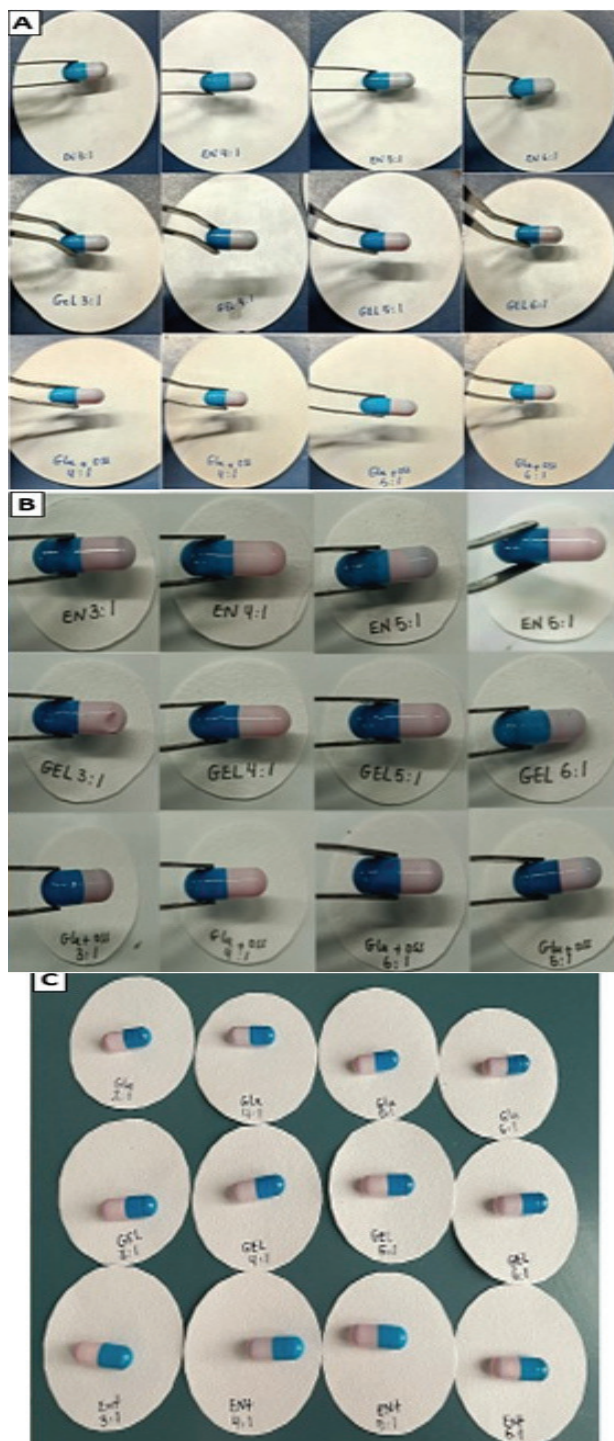


Fig. S-7. Photographs of enteric (Ent), gelatin (Gel) and glutinous capsules taken after 24 h filled with the tested DESs of different molar ratios at (A) 25 °C, (B) 30 °C, and (C) 40 °C.

REFERENCES

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