

Supplementary Material for:

Rapid Polymorphic Screening using Sessile Microdroplets: Competitive Nucleation of Mannitol Polymorphs

Ruel Cedeno¹, Romain Grossier¹, Nadine Candoni¹, Stéphane Veessler^{1*}

¹CNRS, Aix-Marseille Université, CINaM (Centre Interdisciplinaire de Nanosciences de Marseille), Campus de Luminy, Case 913, F-13288 Marseille Cedex 09, France

S1. Evaporation Model

Table S1 Numerical values used as input in evaporation model¹ for aqueous D-mannitol droplets and Mersmann correlation², referenced at 25°C and 1 atm.

Quantity	Symbol	Value	Unit
solubility of water in PDMS oil ¹	c_s	8.76	mol/m ³
diffusivity of water in PDMS oil ¹	D	6.74×10^{-9}	m ² s ⁻¹
supersaturation at matching time*	S_{match}	4.00	-
coefficient of density change for mannitol ³	b_1	0.0719	-
coefficient of water activity lowering for mannitol ⁴	b_2	0.194	-
solubility of β -mannitol in water ⁵	c_{eq}	1.185	mol/kg
solubility of β -mannitol in water ⁵	c_{eq}	1217.4	mol/m ³
solubility of α -mannitol in water ⁵	c_{eq}	1453.0	mol/m ³
solubility of δ -mannitol in water ⁵	c_{eq}	1667.5	mol/m ³
equilibrium solid-state β -mannitol ⁵	C_{sol}	8323.7	mol/m ³
equilibrium solid-state α -mannitol ⁵	C_{sol}	8210.5	mol/m ³
equilibrium solid-state δ -mannitol ⁵	C_{sol}	8365.6	mol/m ³
molar mass of mannitol	M_{Man}	0.182	kg/mol
diffusivity of mannitol in water ⁶	D_i	6.05×10^{-10}	m ² /s
density of pure water ⁷	ρ_w	997	kg/m ³

*measured by monitoring/interpolating the droplet volume (lateral view) as it optically disappears, supersaturation $S = c/c_{eq}$

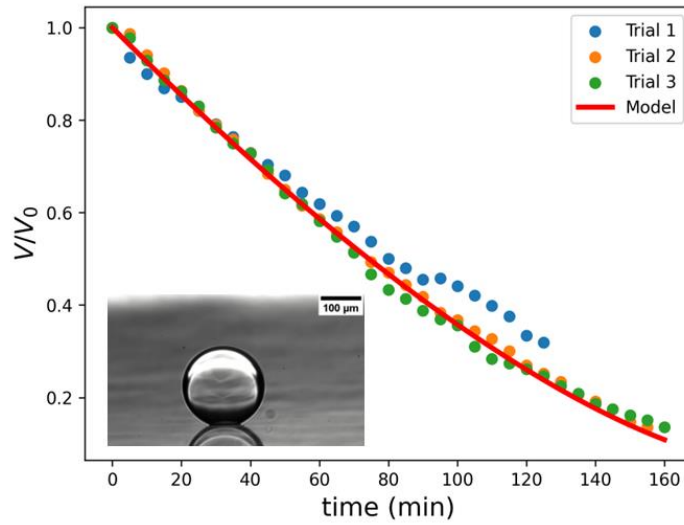


Figure S1. Evolution of relative droplet volume (V/V_0) as a function of time (3 replicates) taken from lateral images.

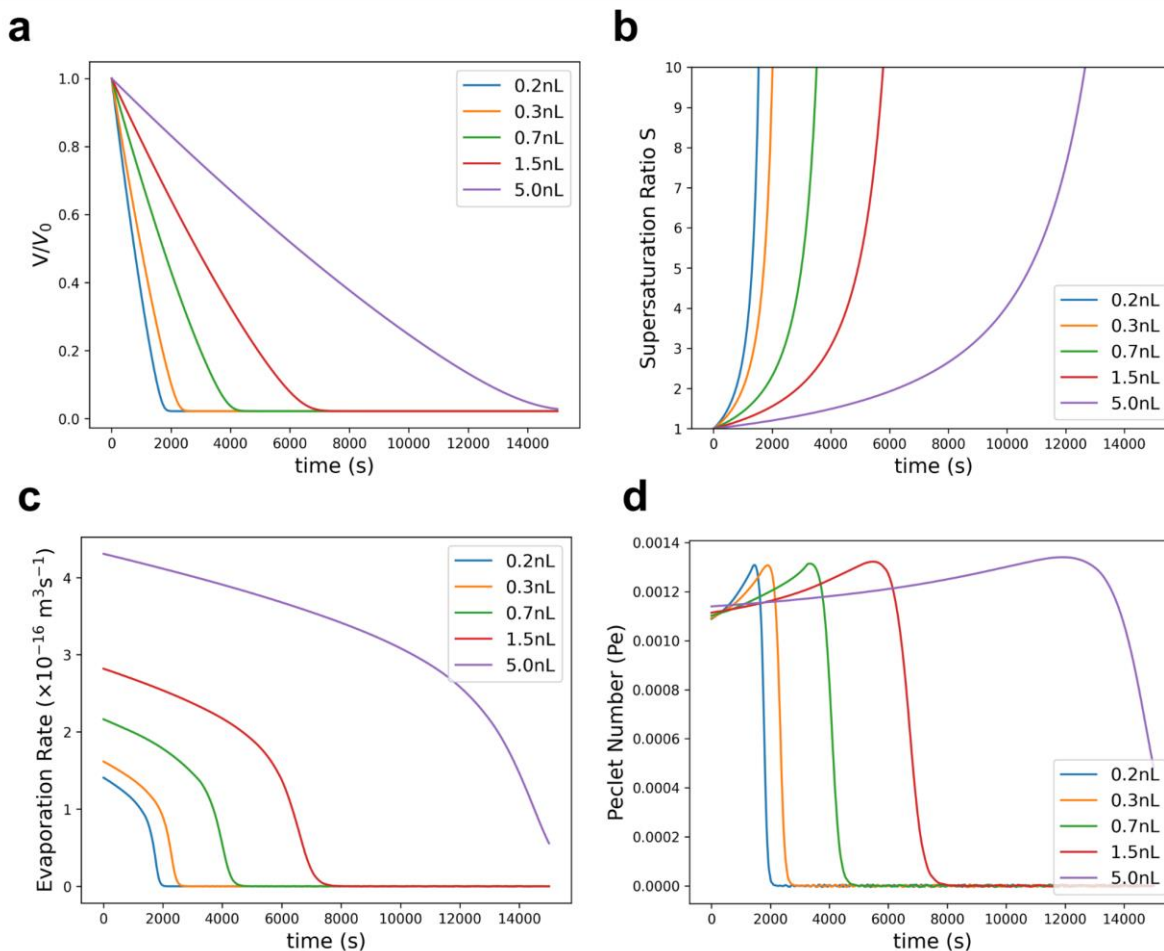


Figure S2. Model predictions for bottom-view arrays of microdroplets in terms of (a) relative volume V/V_0 (b) supersaturation ratio (c) evaporation rate (d) Peclet number^{1, 8}. $Pe < 1$ suggests a uniform distribution of concentration within the droplet. The droplets were subjected under the following conditions: $RH = 0.10$, $T = 25^\circ\text{C}$, $P = 1 \text{ atm}$.

REFERENCES

1. Cedeno, R.; Grossier, R.; Tishkova, V.; Candoni, N.; Flood, A. E.; Veessler, S., Evaporation Dynamics of Sessile Saline Microdroplets in Oil. *Langmuir* **2022**, *38* (31), 9686-9696.
2. Mersmann, A., Calculation of interfacial tensions. *J. Cryst. Growth* **1990**, *102* (4), 841-847.
3. Zhu, C.; Ma, Y.; Zhou, C., Densities and Viscosities of Sugar Alcohol Aqueous Solutions. *Journal of Chemical & Engineering Data* **2010**, *55* (9), 3882-3885.
4. Ninni, L.; Camargo, M. S.; Meirelles, A. J. A., Water Activity in Polyol Systems. *Journal of Chemical & Engineering Data* **2000**, *45* (4), 654-660.
5. Su, W.; Hao, H.; Glennon, B.; Barrett, M., Spontaneous Polymorphic Nucleation of d-Mannitol in Aqueous Solution Monitored with Raman Spectroscopy and FBRM. *Cryst. Growth Des.* **2013**, *13* (12), 5179-5187.
6. Bashkatov, A. N.; Genina, E. A.; Sinichkin, Y. P.; Kochubey, V. I.; Lakodina, N. A.; Tuchin, V. V., Glucose and mannitol diffusion in human dura mater. *Biophys. J.* **2003**, *85* (5), 3310-8.
7. Don, W. G.; Robert, H. P., *Perry's Chemical Engineers' Handbook, Eighth Edition*. 8th ed. / ed.; McGraw-Hill Education: New York, 2008.
8. Gregson, F. K. A.; Robinson, J. F.; Miles, R. E. H.; Royall, C. P.; Reid, J. P., Drying Kinetics of Salt Solution Droplets: Water Evaporation Rates and Crystallization. *The Journal of Physical Chemistry B* **2019**, *123* (1), 266-276.