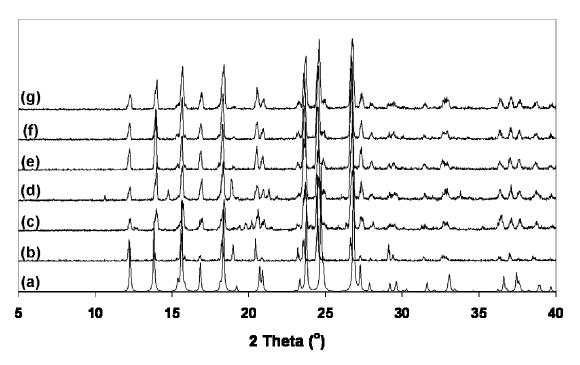
## **Supporting Information**

S1. X-ray data for acetaminophen crystallized from pure ethanol and in the presence of  $\alpha$ -lactose monohydrate, D-mannitol, graphite, L-histidine and sodium chloride.



**Figure S1.** X-ray diffraction data for (a) AAP Form I simulated in Mercury 2.4 (b) AAP from pure ethanol (c) AAP crystallized in the presence of α-lactose monohydrate (d) AAP crystallized in the presence of D-mannitol (e) AAP crystallized in the presence of graphite (f) AAP crystallized in the presence of L-histidine and (g) AAP crystallized in the presence of sodium chloride.

Diffraction pattern simulated in Mercury 2.4 using CIF file found in Cambridge Structural Database (REF Code: HXACAN07).

## S2. Optical micrographs of the $\alpha\text{-lactose}$ monohydrate and D-mannitol crystals used in the induction time measurements.

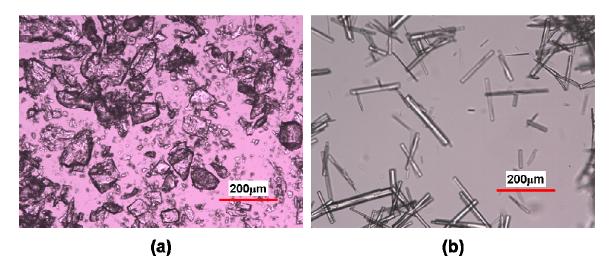
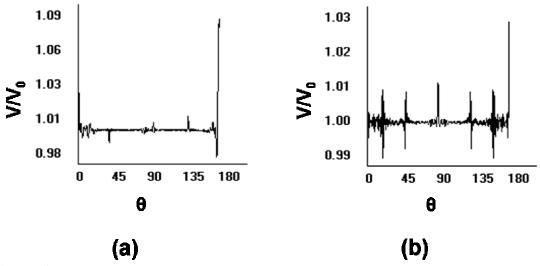


Figure S2. Optical micrographs of (a)  $\alpha$ -lactose monohydrate and (b) D-mannitol crystals used in the acetaminophen induction time experiments.

S3. EpiCalc epitaxy calculation for (0-11) Form I acetaminophen with (0-11)  $\alpha$ -lactose monohydrate and (100) Form I acetaminophen with (00-1) D-mannitol.



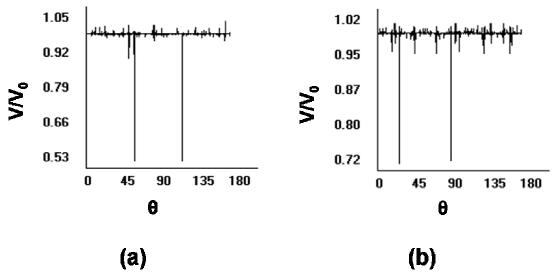
**Figure S3.** A plot of dimensionless potential energy  $(V/V_0)$  as a function of orientation angle (θ) for **a**) (0-11) face of acetaminophen and (0-11) face of α-lactose monohydrate and **b**) (100) face of acetaminophen and (00-1) of D-mannitol.

The EpiCalc calculations were performed using the following lattice values for the (0-11) and (100) faces of Form I acetaminophen, the (0-11) face of  $\alpha$ -lactose monohydrate and the (00-1) face of D-mannitol:

- (0-11) Form I acetaminophen:  $a = 14.84 \text{ Å}, b = 7.09 \text{ Å} \text{ and } \beta = 96.12^{\circ}$
- (100) Form I acetaminophen:  $a = 11.62 \text{ Å}, b = 9.23 \text{ Å} \text{ and } \beta = 90^{\circ}$
- (0-11)  $\alpha$ -lactose monohydrate: a = 15.19 Å, b = 9.57 Å and  $\alpha = 95.85^{\circ}$
- (00-1) D-mannitol:  $a = 16.9 \text{ Å}, b = 8.69 \text{ Å} \text{ and } \alpha = 90^{\circ}$

The acetaminophen overlayer dimensions were 25 x 25 cells and the orientation angle range tested was  $180^{\circ}$  with a step size of  $0.25^{\circ}$ .

S4. EpiCalc epitaxy calculation for the (100) and (0-11) faces of Form I acetaminophen with the (002) face of graphite



**Figure S4.** A plot of dimensionless potential energy  $(V/V_0)$  as a function of orientation angle  $(\theta)$  for **a**) (100) face of acetaminophen and (002) face of graphite and **b**) (0-11) face of acetaminophen and (002) of graphite.

The EpiCalc calculations were performed using the following lattice values for the (100) and (0-11) faces of Form I acetaminophen and the (002) face of graphite:

(100) Form I acetaminophen: a = 11.62 Å, b = 9.23 Å and  $\beta$  =  $90^{\rm o}$ 

(0-11) Form I acetaminophen:  $a = 14.84 \text{ Å}, b = 7.09 \text{ Å} \text{ and } \beta = 96.12^{\circ}$ 

(002) graphite: a=b=2.456~Å and  $\alpha=120^{o}$ 

The acetaminophen overlayer dimensions were 25 x 25 cells and the orientation angle range tested was  $180^{\circ}$  with a step size of  $0.25^{\circ}$ .

S5. Table displaying the minimum interaction energy and optimal orientation angle between the D-mannitol (00-1) face and acetaminophen (100) face at different interplanar distances

Azimuthal Angle (°)	Interaction energy per cell (kcal/mol)
90	-0.50
	-0.55
	-0.61
270	-0.66
270	-0.72
270	-0.78
270	-0.83
90	-0.92
90	-1.01
90	-1.11
90	-1.24
90	-1.39
90	-1.56
90	-1.72
75	-1.91
25	-2.16
25	-2.45
25	-2.77
25	-3.06
270	-3.33
90	-3.78
90	-4.28
90	-4.78
90	-4.99
90	-4.85
155	-4.84
270	-5.15
270	-5.59
270	-5.61
270	-4.53
260	-1.93
260	3.94
260	20.93
90	57.26
90	108.45
	Angle (°)  90 270 270 270 270 270 270 90 90 90 90 90 90 75 25 25 25 25 25 270 90 90 90 90 90 270 270 270 270 270 270 270 270 270 27

S6. Table displaying the minimum interaction energy and optimal orientation angle between the  $\alpha$ -lactose (0-11) face and acetaminophen (0-11) face at different interplanar distances

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Interplanar distance (Å)	Azimuthal Angel (°)	Interaction energy per cell (kcal/mol)
6	265	-2.00
5.8	265	-2.25
5.6	265	-2.53
5.4	265	-2.84
5.2	265	-3.17
5	265	-3.56
4.8	265	-4.01
4.6	265	-4.53
4.4	265	-5.08
4.2	265	-5.61
4	265	-6.33
3.8	85	-7.20
3.6	85	-8.20
3.4	85	-9.19
3.2	85	-10.21
3	85	-11.31
2.8	85	-12.21
2.6	85	-13.33
2.4	85	-13.84
2.2	85	-12.98
2	265	-10.48
1.8	265	-8.39
1.6	265	-2.10
1.4	265	12.98
1.2	265	45.84
1	265	114.05
0.8	265	247.20
0.6	265	547.61
0.4	265	1200.92
0.2	265	3233.64