

SUPPLEMENTARY INFORMATION

A co-crystalline solid solution affords a high-soluble and fast-absorbing form of praziquantel

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Reaction Summary

Table S1: Summary of the product identified in this work

R-PZQ	S-PZQ	D-MA	L-MA	D-TA	L-TA	Product
0.5	0.5	0	1	0	0	CC1; CC2
1.5	1.5	1	2	0	0	SS3
1	1	1	1	0	0	CC3
0.5	0.5	0	0	0	1	CC5
1.5	1.5	0	0	1	2	SS4
1	1	0	0	1	1	CC4
1.5	1.5	0	2	1	0	CC1; CC2; 5
1	1	0	1	1	0	SS5
1.5	1.5	0	1	2	0	SS5
1.5	1.5	0	2	0	1	CC1; CC2; 5
1	1	0	1	0	1	SS5
1.5	1.5	0	1	0	2	SS5
2	2	1	1	0	2	SS5
2	2	0	1	1	2	SS4
2	2	1	1	1	1	CC3 CC4

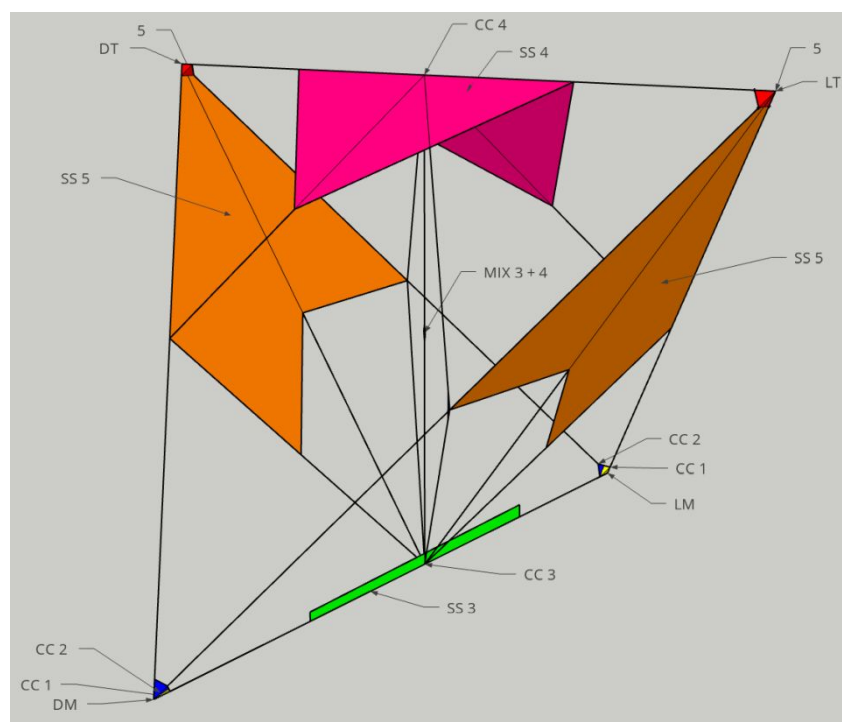


Figure S1. 3-D phase diagram for PZQ in combination with four dicarboxylic acids.

Crystallographic information

Table S1: Crystal data and structure refinement details.

	R-PZQ:S-PZQ:D-MA:L-MA (3)	R-PZQ:S-PZQ:D-TA:L-TA (4)
Empirical formula	C ₂₃ H ₂₇ N ₂ O ₇	C ₂₃ H ₂₈ N ₂ O ₈
Formula weight (g mol ⁻¹)	443.473	460.480
T (K)	293	293
Wavelength (Å)	1.540598	1.540598
Crystal system	orthorhombic	triclinic
Space group	P b c a	P-1
<i>a</i> (Å)	20.150(4)	12.561(3)
<i>b</i> (Å)	25.108(5)	10.749(2)
<i>c</i> (Å)	8.8963(16)	9.2445(17)
α (°)	90	110.316(4)
β (°)	90	95.675(4)
γ (°)	90	100.004(6)
<i>V</i> (Å ³)	4501.0(15)	1135.5(4)
<i>Z</i>	8	2
ϑ range for data collection (°)	5° to 60°	5° to 60°
<i>R</i> _{int}	0.07524	0.06217
Refinement method	Rietveld	Rietveld
Number parameters	34	181
Goodness-of-fit	2.70384	2.40215
<i>R</i> _{wp}	0.10378	0.08239
<i>R</i> _{exp}	0.03838	0.03430

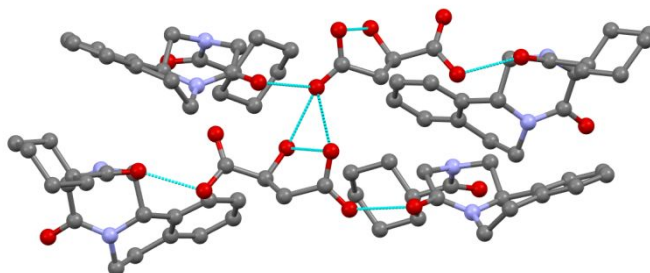


Figure S2: H-bond motif across multiple chains in R-PZQ:S-PZQ:D-MA:L-MA (3).

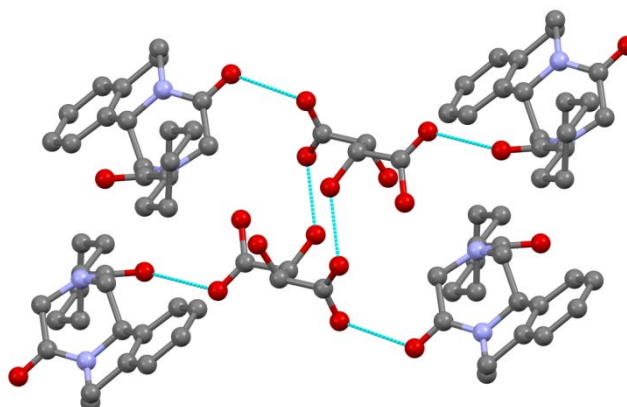


Figure S3: H-bond motif across multiple chains in R-PZQ:S-PZQ:D-TA:L-TA (4).

Rietveld refinement

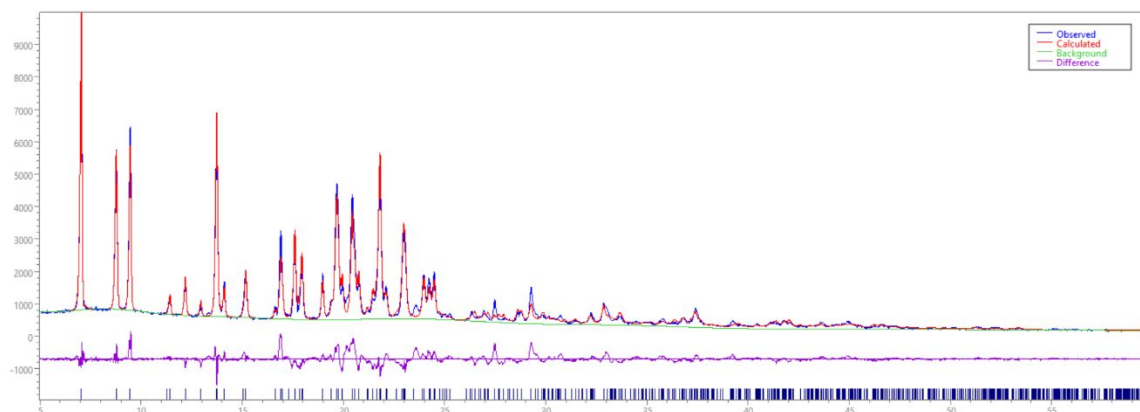


Figure S4: Rietveld refinement (red line) of R-PZQ:S-PZQ:D-MA:L-MA (**3**) compared with the experimental diffraction pattern (blue line). The purple line indicates the difference plot.

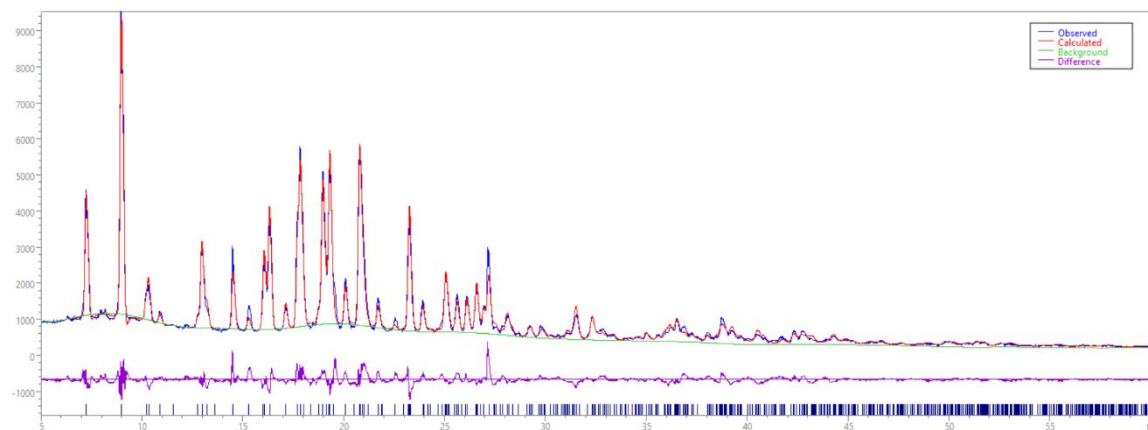


Figure S5: Rietveld refinement (red line) of R-PZQ:S-PZQ:D-TA:L-TA (**4**) compared with the experimental diffraction pattern (blue line). The purple line indicates the difference plot.

5-component ball milling

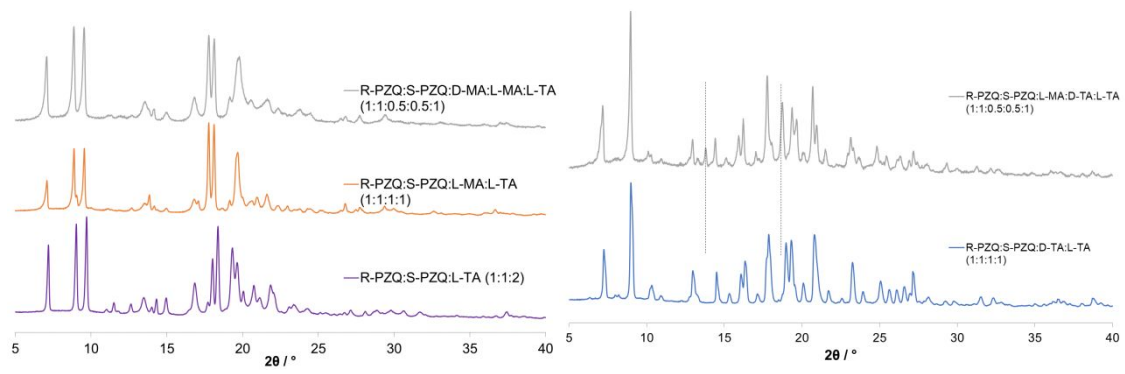


Figure S6: PXRD patterns for the 5-component solid solutions

Assessment of solution-mediated transformations

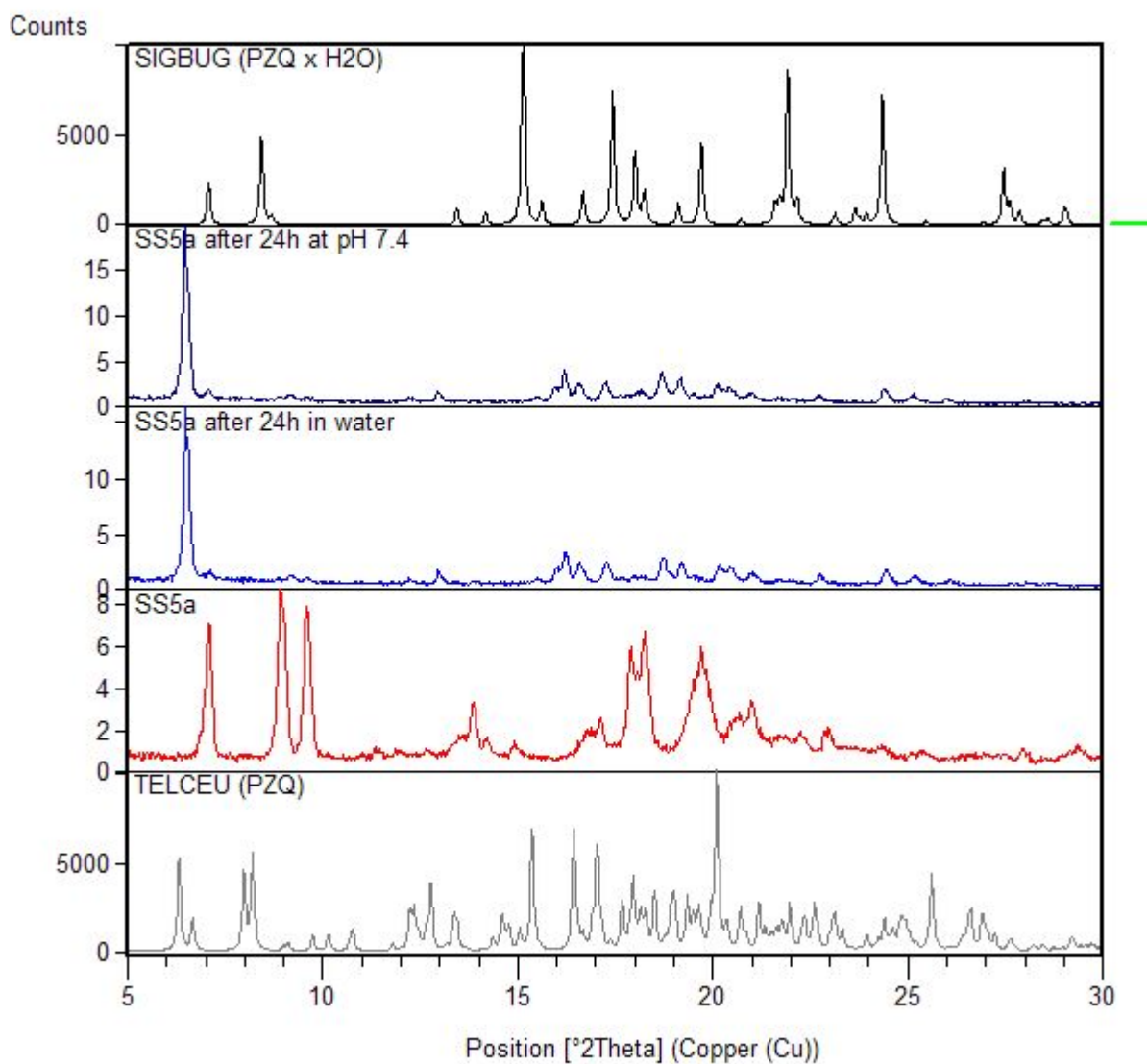


Figure S7 Comparison of PXR patterns calculated for pure PZQ and PZQxH₂O against those measured for SS5a as synthesised, after 24h in H₂O and after 24h in a buffer solution at pH 7.4

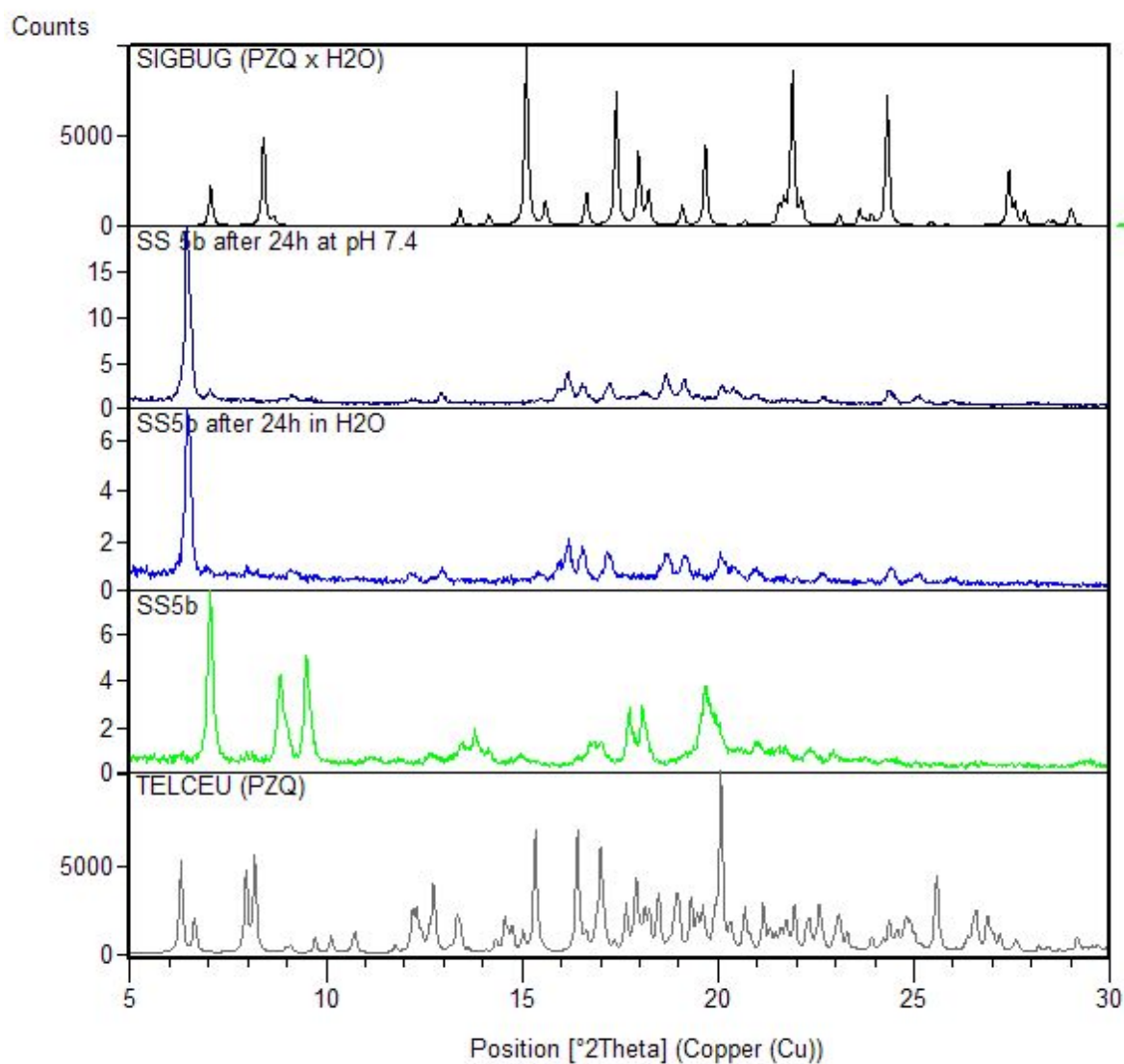


Figure S8 Comparison of PXRD patterns calculated for pure PZQ and PZQxH₂O against those measured for SS5a as synthesised, after 24h in H₂O and after 24h in a buffer solution at pH 7.4