

**Supplementary Table S1: Molecular weight and hydrodynamic radius of Xanthan, Alginate and IDeg**

|          | Molecular Weight (kDa) | Hydrodynamic radius* <sup>2</sup> at 20° | Hydrodynamic radius* <sup>2</sup> at 37° |
|----------|------------------------|--|--|
| Xanthan  | 261                    | 79                                       | 75                                       |
| Alginate | 56                     | 15                                       | 15                                       |
| IDeg     | 73* <sup>1</sup>       | 5  | 4.4                                      |

\*<sup>1</sup>- Molecular weight of a dodecamer of insulin

\*<sup>2</sup> -Calculated using  $R_h = \left( \frac{3[\eta]M}{10\pi N} \right)^{1/3}$

Armstrong, J.K., et al., *The hydrodynamic radii of macromolecules and their effect on red blood cell aggregation*. Biophysical journal, 2004. 87(6): p. 4259-4270.

**Supplementary Table S2:** Intrinsic viscosity (ml/g) of insulin degludec (IDeg), xanthan (X), alginate (A), Binary system containing xanthan and alginate (XA) and PIC measured at 20°C and 37°C

|             | Intrinsic viscosity at 20°C | R <sup>2</sup> | Intrinsic viscosity at 37°C | R <sup>2</sup> | P value               |
|-------------|-----------------------------|----------------|-----------------------------|----------------|-----------------------|
| <b>X</b>    | 12000 ± 520                 | 0.987          | 10300 ± 6000                | 0.968          | 0.55                  |
| <b>A</b>    | 360 ± 31                    | 0.996          | 360±11                      | 0.999          | 0.41                  |
| <b>IDeg</b> | 9.2 ± 1.4                   | 0.950          | 7.4 ±1.8                    | 0.947          | 0.022                 |
| <b>XA</b>   | 1000 ± 160                  | 0.976          | 590 ± 60                    | 0.996          | 3.8x10 <sup>-13</sup> |
| <b>PIC</b>  | 490 ± 60                    | 0.998          | 710 ± 58                    | 0.998          | 1.2x10 <sup>-13</sup> |

**Supplementary Table S3:** Results for ANOVA test for PSD of Xanthan (X), alginate (A), Insulin degludec (IDeg), XA, and PIC at a range of temperatures (4, 18 and 37°C) on Day 1, Day 7, and Day 14

|             | Day 1            | Day 7    | Day 14            |          |                  |          |
|-------------|------------------|----------|-------------------|----------|------------------|----------|
| <b>X</b>    | $F_{1,2}=33.50$  | $P=0.11$ | $F_{1,2}=35.49$   | $P=0.11$ | $F_{1,2}=301.42$ | $P=0.04$ |
| <b>A</b>    | $F_{1,2}=28.42$  | $P=0.12$ | $F_{1,2}=60.45$   | $P=0.08$ | $F_{1,2}=14.31$  | $P=0.16$ |
| <b>IDeg</b> | $F_{1,2}=375.01$ | $P=0.03$ | $F_{1,2}=25.17$   | $P=0.13$ | $F_{1,2}=1.16$   | $P=0.48$ |
| <b>XA</b>   | $F_{1,2}= 2.44$  | $P=0.36$ | $F_{1,2}=4382.43$ | $P=0.01$ | $F_{1,2}=4.52$   | $P=0.28$ |
| <b>PIC</b>  | $F_{1,2}=1.07$   | $P=0.49$ | $F_{1,2}=3395.93$ | $P=0.01$ | $F_{1,2}=0.30$   | $P=0.68$ |

**Supplementary Table S4:** MANOVA analysis

| Component                               | F statistic       | Significance |
|---|-------------------|--------------|
| <b>Sample</b>                           | $F_{4,1765}=6543$ | $P<0.001$    |
| <b>Concentration</b>                    | $F_{10,1765}=487$ | $P<0.001$    |
| <b>Temperature</b>                      | $F_{7,1765}=1181$ | $P<0.001$    |
| <b>Sample*Concentration</b>             | $F_{39,1765}=354$ | $P<0.001$    |
| <b>Sample*Temperature</b>               | $F_{28,1765}=51$  | $P<0.001$    |
| <b>Temperature*Concentration</b>        | $F_{70,1765}=13$  | $P<0.001$    |
| <b>Sample*Temperature*Concentration</b> | $F_{273,1765}=4$  | $P<0.001$    |