**Figure SI**: Morphology of microparticles prepared by spray drying and observed by digital optical microscopy (X40).(A&B): Optical Micrographs of DEC & DOX Mps Uncrosslinked (C& D) Microscopic images of Chi-DEC MPs & Chi-DOX

В D6: 1.081016 n D16: 9 microns D10: 1.893921 mics : 1.6112 rons n, 6 1.174560 microns 236 microns

| Batch No.   | D[v, 10] μm      | D[v, 50] μm | D[v, 90] μm | Zeta<br>Potential(mV) |
|-------------|------------------|-------------|-------------|-----------------------|
| DEC MPs     | $3.552 \pm 0.61$ | 4.412±0.78  | 5.688±0.54  | -28.8± 0.79           |
| Chi-DEC MPs | 3.549±0.65       | 4.605±0.59  | 5.690±0.55  | +27.1±0.85            |
| DOX MPs     | $3.655 \pm 0.68$ | 4.418±0.56  | 5.677±0.56  | -25.8± 0.88           |
| Chi-DOX MPs | $3.055 \pm 0.75$ | 4.740± 0.56 | 7.020± 0.59 | +29.6±0.85            |
| Blank MPs   | 3.461± 0.80      | 4.588± 0.91 | 5.719± 0.65 | -31.9±0.55            |

Table S1. Particle size distribution and Zeta potential of spray-dried microparticles determined by laser diffraction (n = 3), expressed as volume mean diameter, D ( $\mu$ m).

Data represented as mean  $\pm$  S.D (n=3). D[v, 10] ;diameter is the diameter at which 10% of a sample's mass is comprised of smaller particles. D [v, 50]; is the diameter at which 50% of a sample's mass is comprised of smaller particles. D[v, 90] is the diameter at which 90% of a sample's mass is comprised of smaller particles

 Table S2. Size, zeta potential and Drug entrapement of formulation stability conditions at

 different temperature conditions after Six months.

| Formulation    | Temperatu<br>re(°C)&<br>Relative<br>Humidity<br>% | Drug Entra | apment (%)        | Particle<br>(µ | size ± SD<br>um)  | Zeta potential<br>(mV) |                   |
|----------------|---|------------|-------------------|----------------|-------------------|------------------------|-------------------|
|                |   | Initial    | After 6<br>months | Initial        | After 6<br>months | Initial                | After 6<br>months |
|                |   |            |                   |                |                   | +27.6±                 | +27.8±            |
| CHI-DEC<br>MPs | 2-8°C   | 41.28±1.5  | 41.11±1.5         | 4.81±0.5       | 4.89±0.59         | 0.28                   | 0.59              |
| IVII S         | 25 °C / 60%                                       |            |                   |                |                   | +27.7±                 | +27.9±            |
|                | RH  | 41.56±1.7  | 41.22±1.6         | 4.83±0.6       | 4.83±0.68         | 0.37                   | 0.61              |
|                | 40°C / 75%  |            |                   |                |                   | +27.5±                 | +27.3±            |
|                | RH  | 40.56±1.8  | 39.86±1.8         | 4.83±0.8       | 4.83±0.81         | 0.63                   | 0.79              |
|                |   |            |                   |                |                   | +29.4±                 | +29.6±            |
|                | 2-8°C   | 43.51±1.9  | 43.0±1.7          | 4.79±0.7       | 4.79±0.78         | 0.41                   | 0.49              |
| Chi-DOX        | 25 °C / 60%                                       |            |                   |                |                   | +29.7±                 | +29.5±            |
| MPs            | RH  | 44.62±1.8  | 44.32±1.9         | 4.93±0.8       | 4.93±0.83         | 0.49                   | 0.51              |
|                | 40°C / 75%  |            |                   |                |                   | +28.9±                 | +29.2±            |
|                | RH  | 42.51±2.0  | 40.91±1.9         | 5.3±0.8        | 5.3±0.88          | 0.69                   | 0.87              |

| Table S3. | In vitro antifilarial        | activity of Formul     | ated DEC, DOX a    | nd DEC+ DOX on |
|-----------|------------------------------|------------------------|--------------------|----------------|
| :         | adult <i>B. malayi</i> and r | nicrofilariae , deteri | nined after 48 hrs |                |

| Dosogo                        | Activity on Adult   |                               |                               |        | Activity on mf   |                    |                 |
|-------------------------------|---------------------|-------------------------------|-------------------------------|--------|------------------|--------------------|-----------------|
| Dosage                        | MIC <sup>#</sup>    | IC <sub>50</sub> <sup>#</sup> | CC <sub>50</sub> <sup>#</sup> | SI     | MIC <sup>#</sup> | IC <sub>50</sub> # | SI <sup>#</sup> |
| Chi-DEC MPs                   | 25                  | 4.2                           | >100                          | >23.8  | 25               | 3.5                | >28.57          |
| Chi-DOX MPs                   | 10                  | 3.75                          | >100                          | >26.66 | 10               | 2.0                | >45             |
| Chi-DEC MPs + Chi-<br>DOX MPs | (25 DEC<br>+10 DOX) | 2.10                          | >100                          | >47.62 | 2.5              | 1.9                | >55.63          |

#concentration in  $\mu M$ 

Table S4. *In vitro* motility scoring and % inhibition in MTT reduction results of Formulated DEC, DOX and DEC+ DOX on adult *B. malayi* and microfilariae (n=3), was determined at 24 h.

| Drugs        | Conc. µM Adult worm         |          |              | Motility     |  |
|--------------|-----------------------------|----------|--------------|--------------|--|
|              |                             | Motility | % inhibition | - scoring Mf |  |
|              |                             | scoring  | reduction    |              |  |
| Chi-DEC MPs  | 25                          | D        | 70.2±2.8     | D            |  |
|              | 10                          | 1+       | 41.5±1.6     | 1+           |  |
|              | 5                           | 1+       | 30.4±3.2     | 1+           |  |
|              | 2.5                         | 2+       | 12.2±2.4     | 1+           |  |
|              | 1.25                        | 2+       | 10.5±2.1     | 2+           |  |
| Chi-DOX MPs  | 10                          | D        | 65.5±3.8     | D            |  |
|              | 5                           | 1+       | 50±4.2       | D            |  |
|              | 2.5                         | 1+       | 20.6±1.8     | 1+           |  |
|              | 1.25                        | 2+       | 18.9±3.6     | 2+           |  |
|              | 0.62                        | 2+       | 18±1.2       | 2+           |  |
| Chi-DEC MPs+ | 25 μMDEC +10μM DOX          | D        | 82.5±3.2     | D            |  |
| Chi-DOX MPs  | 10 μMDEC + 5 μMDOX          | 1+       | 52±1.8       | D            |  |
|              | $5 \mu MDEC + 2.5 \mu MDOX$ | 2+       | 40.9±3.6     | 1+           |  |
|              | 2.5µMDEC+1.25µMDOX          | 2+       | 22±1.4       | 1+           |  |
|              | 1.25µMDEC +0.6µMDOX         | 2+       | 14±2.8       | 2+           |  |

#Scored as 0% motility reduction (4+); 1–49% motility reduction (3+); 50–74% motility reduction (2+); 75–99% motility reduction (1+) and D represents 100% motility reduction (Dead).

| Table S5. R <sup>2</sup> | <sup>2</sup> value of different | t release kinetio | c models |
|--------------------------|---------------------------------|-------------------|----------|
|--------------------------|---------------------------------|-------------------|----------|

| Formulation<br>Batch | Zero order<br>R <sup>2</sup> | First<br>order R <sup>2</sup> | Korsmeyer-<br>peppas R <sup>2</sup> | Higuchi<br>R <sup>2</sup> | Hixson crowell<br>R <sup>2</sup> | Weibull<br>R <sup>2</sup> |
|----------------------|------------------------------|-------------------------------|-------------------------------------|---------------------------|----------------------------------|---------------------------|
| Chi-DEC-MPs          | 0.9652                       | 0.9848                        | 0.9864                              | 0.9209                    | 0.9889                           | 0.9900                    |
| Chi-DOX-MPs          | 0.9726                       | 0.9879                        | 0.9907                              | 0.9213                    | 0.9923                           | 0.9941                    |