

**Title:**

*In vitro-in vivo* assessments of apocynin-hybrid nanoparticles-based gel as an effective nanophytomedicine for treatment of rheumatoid arthritis

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**Table S2** Right paw thickness (mm) of all rats from all the experimental groups assessed on days 0, 7<sup>th</sup>, 14<sup>th</sup>, 21<sup>th</sup>, and 28<sup>th</sup>

| Day<br>Group               | 0    | 7    | 14   | 21   | 28   |
|----------------------------|------|------|------|------|------|
| Normal                     | 2.61 | 2.63 | 2.71 | 2.72 | 2.78 |
|                            | 2.45 | 2.51 | 2.89 | 2.59 | 2.77 |
|                            | 2.61 | 2.71 | 2.56 | 2.6  | 2.69 |
|                            | 2.52 | 2.69 | 2.51 | 2.89 | 2.61 |
|                            | 2.5  | 2.48 | 2.52 | 2.46 | 2.55 |
| Plain hybrid NPs-based gel | 2.87 | 5.62 | 5.21 | 5.01 | 4.86 |
|                            | 2.69 | 4.1  | 4.65 | 4.99 | 5.39 |
|                            | 2.94 | 4.9  | 5.1  | 5.78 | 5.28 |
|                            | 2.85 | 4.21 | 5.1  | 5.39 | 4.81 |
|                            | 2.95 | 5.92 | 5.51 | 5.39 | 5.48 |
| APO-hybrid NPs-based gel   | 2.65 | 4.59 | 4.92 | 4.51 | 3.69 |
|                            | 2.69 | 4.29 | 4.65 | 4.64 | 4.39 |
|                            | 2.65 | 6.12 | 5.19 | 4.62 | 4.3  |
|                            | 3.01 | 4.69 | 5.12 | 4.39 | 4.62 |
|                            | 2.36 | 4.19 | 4.25 | 3.56 | 3.54 |
| Olfen <sup>®</sup> gel     | 2.85 | 5.74 | 5.39 | 5.81 | 5.42 |
|                            | 2.48 | 4.51 | 4.67 | 4.59 | 4.51 |
|                            | 3.01 | 5.53 | 5.21 | 6.11 | 5.31 |
|                            | 2.61 | 4.39 | 4.78 | 5.01 | 4.45 |
|                            | 2.74 | 4.19 | 4.81 | 5.08 | 4.73 |
| APO gel                    | 2.76 | 4.99 | 5.22 | 5.71 | 5.31 |
|                            | 2.35 | 4.53 | 4.49 | 4.91 | 4.39 |
|                            | 2.38 | 4.46 | 4.12 | 4.74 | 5.09 |
|                            | 2.88 | 5.29 | 5.12 | 6.53 | 6.21 |
|                            | 2.27 | 4.69 | 4.42 | 4.52 | 4.81 |
| CFA-induced RA             | 2.51 | 5.41 | 5.26 | 7.9  | 7.79 |
|                            | 2.58 | 6.45 | 5.11 | 7.71 | 7.75 |

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|      |      |      |      |      |
|------|------|------|------|------|
| 2.35 | 5.47 | 5.12 | 7.62 | 8.11 |
| 2.51 | 4.89 | 4.75 | 8.22 | 8.46 |
| 3.01 | 5.78 | 5.11 | 7.56 | 7.74 |

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