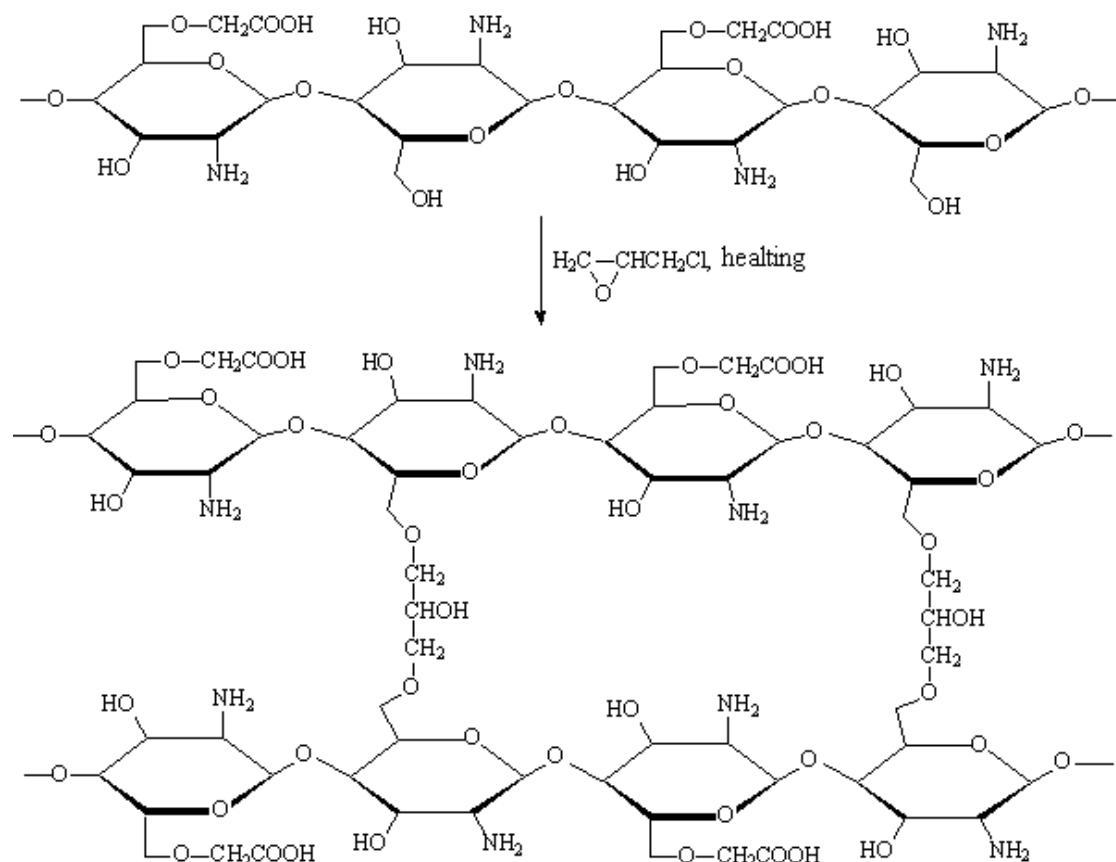


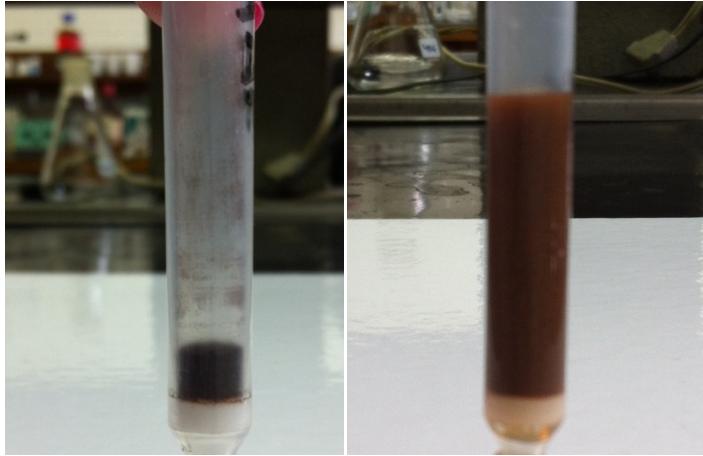
Electronic Supplementary Information (ESI) available

A magnetic nanogel based on chitosan for antitumor drug delivery:

Synthesis, characterization and in vitro drug release.



S1: Crosslinked reaction of O-carboxymethyl chitosan and epichlorohydrin.



a)

b)

S2) Comparison of the swelling the images of the O-CEMg nonogel measured without (a) and with (b) the buffer solution pH 7.4; 100 mg of O-CEMg, contact time 60 min.

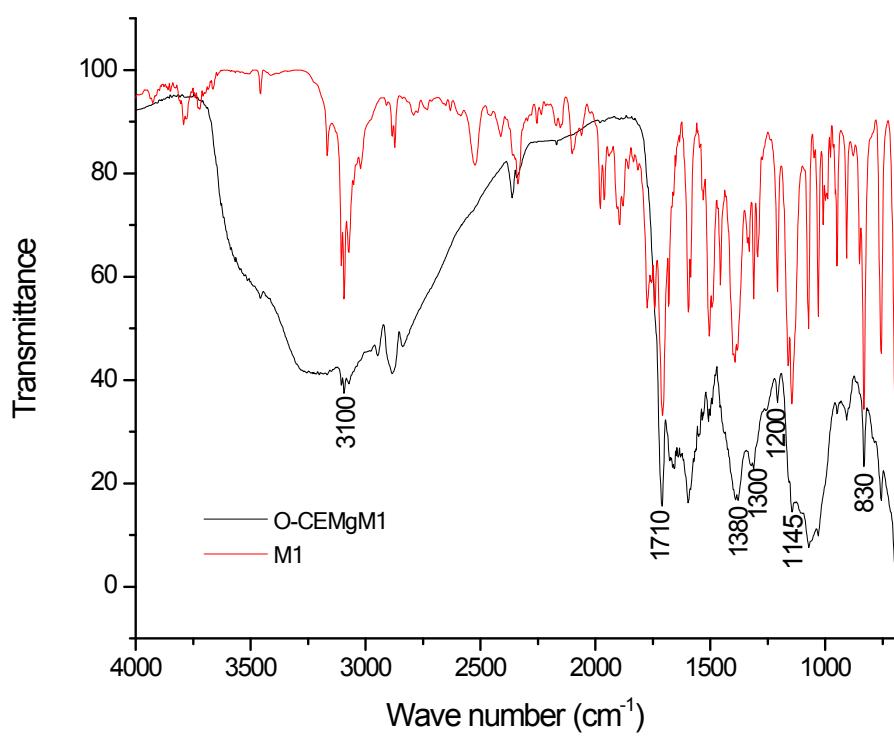


Figure S3: FTIR spectra of M1 loaded in magnetic hydrogel (O-CEMgM1)

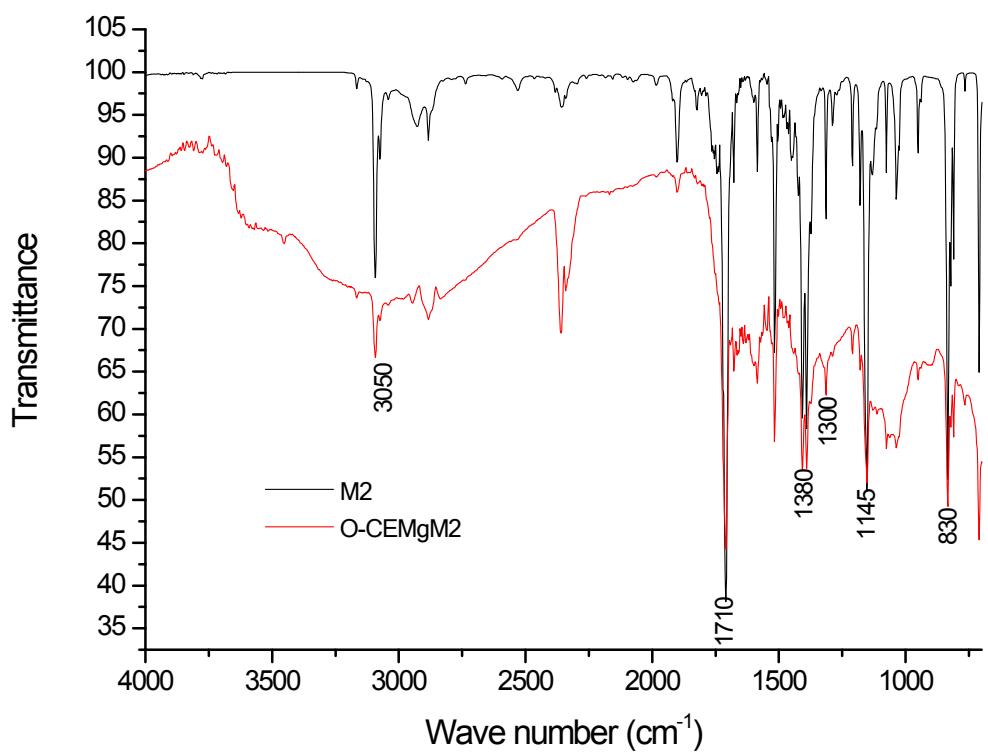


Figure S4: FTIR spectra of M2 loaded in magnetic hydrogel (O-CEMgM2)

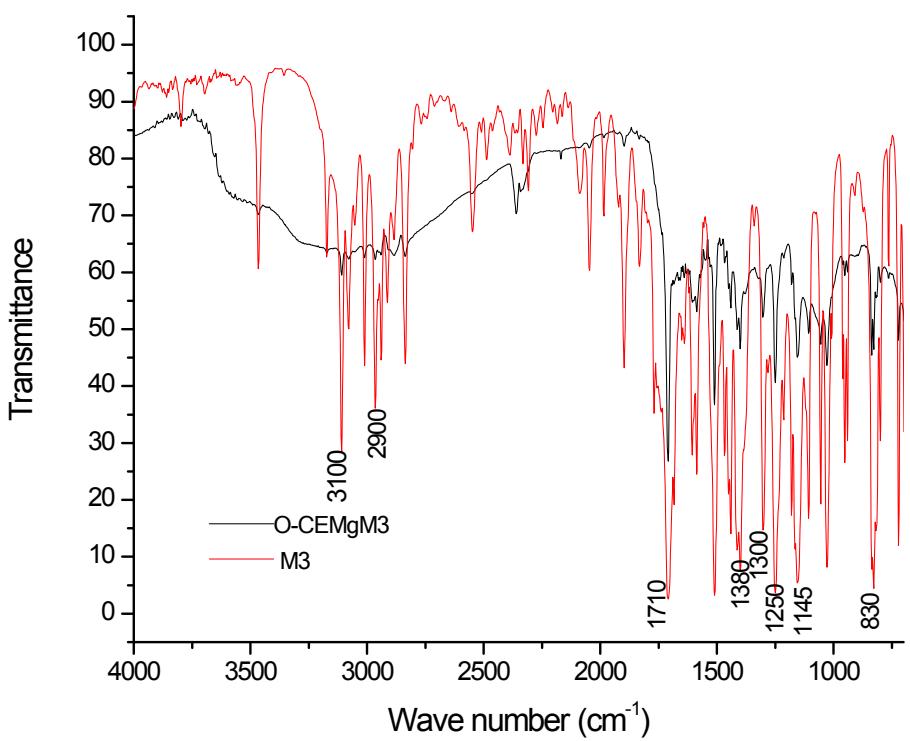


Figure S5: FTIR spectra of M3 loaded in magnetic hydrogel (O-CEMgM3)

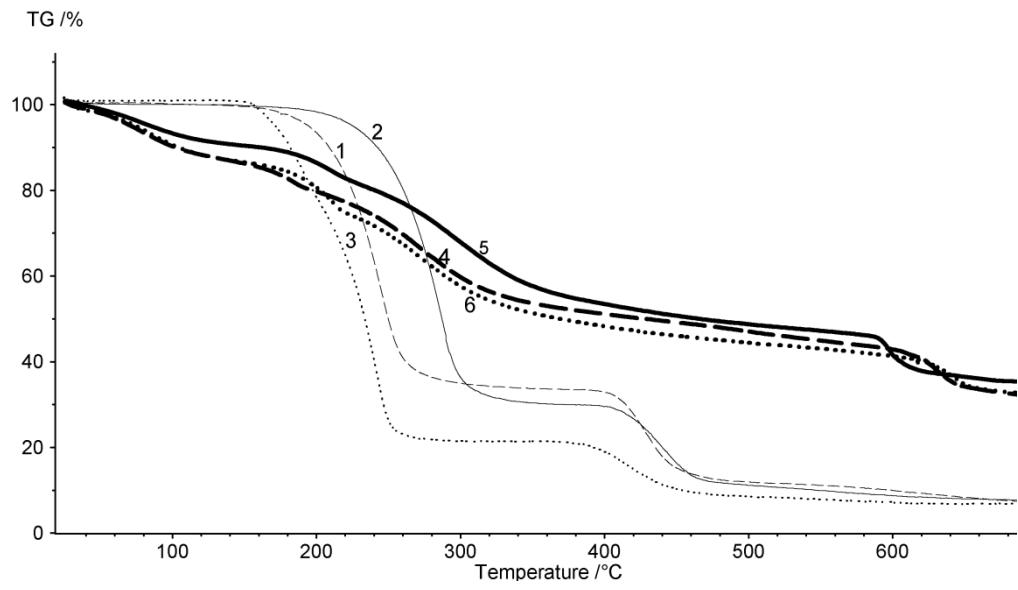


Figure S6: TG curves of M1 (1), M2 (2), M3 (3), O-CEMgM1 (4), O-CEMgM2 (5), O-CEMgM3 (6)

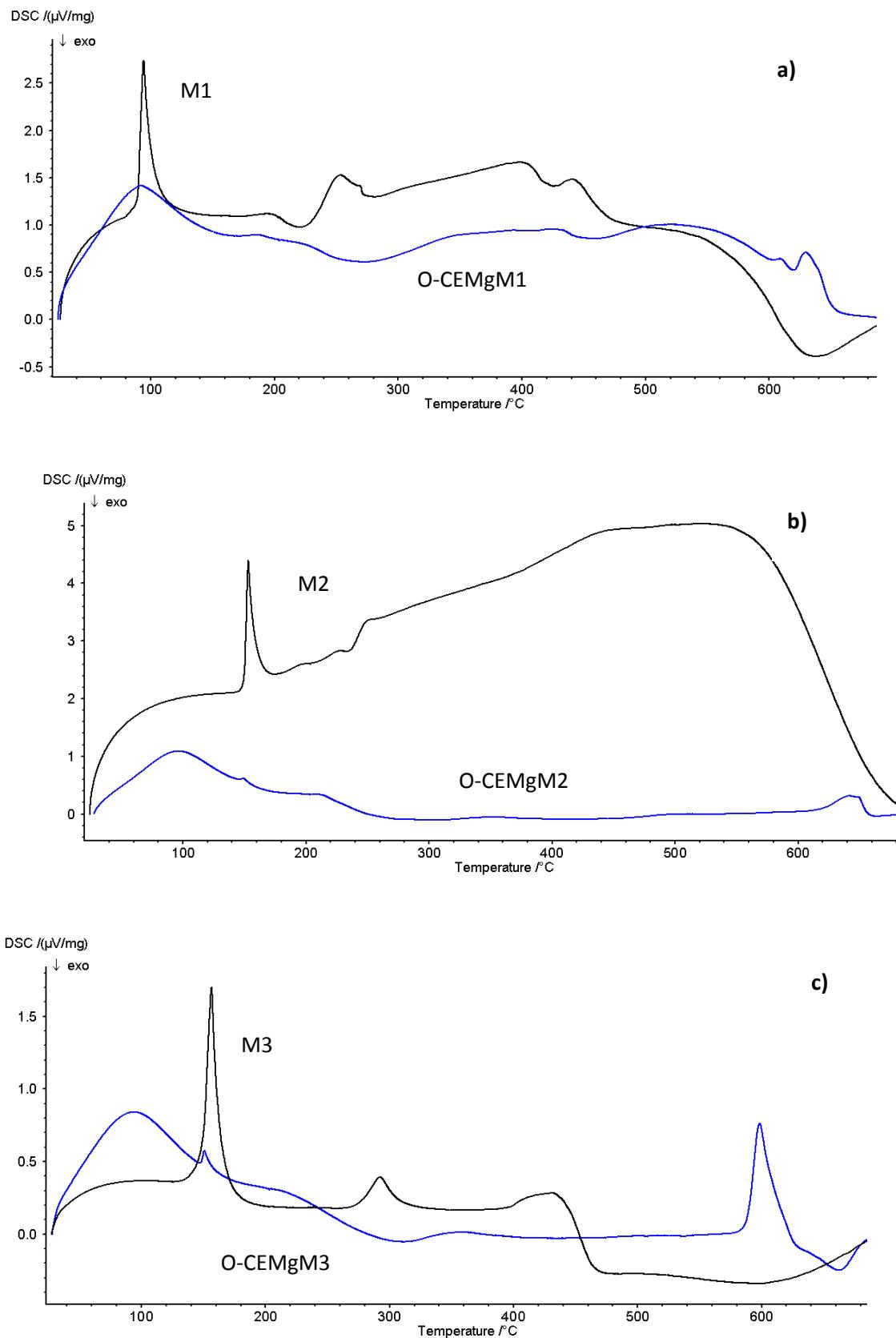


Figure S7: DSC curves of a) M1 and O-CEMgM1; b) M2 and O-CEMgM2; c) M3 and O-CEMgM3.

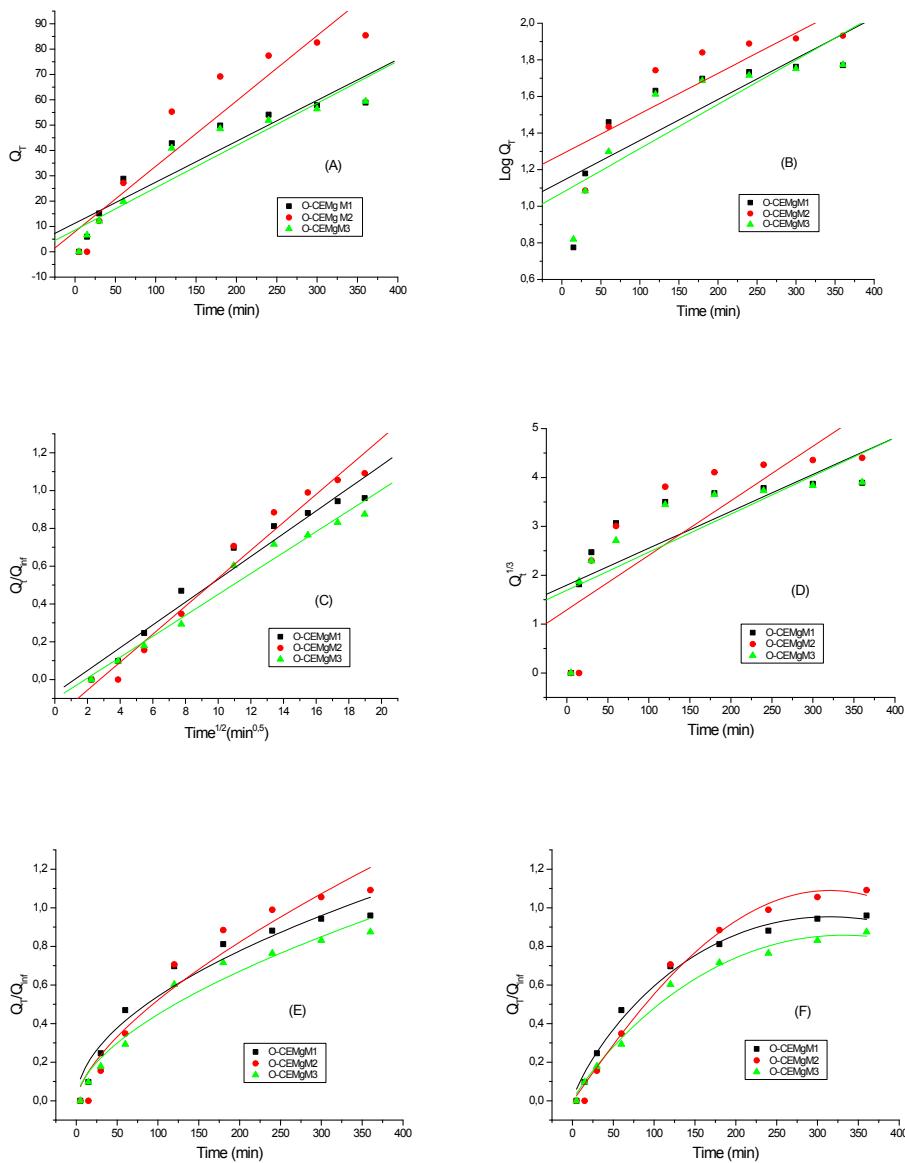


Figure S8: Mathematical models applied to maleimides release in phosphate buffer pH 5.0 to 37 °C of magnetic hydrogel. (A) Order zero, (B) first order (C) Higuchi, (D) Hixson-Crowell, (E)-Peppas and Korsmeyer (F) Peppas-Sahlin.

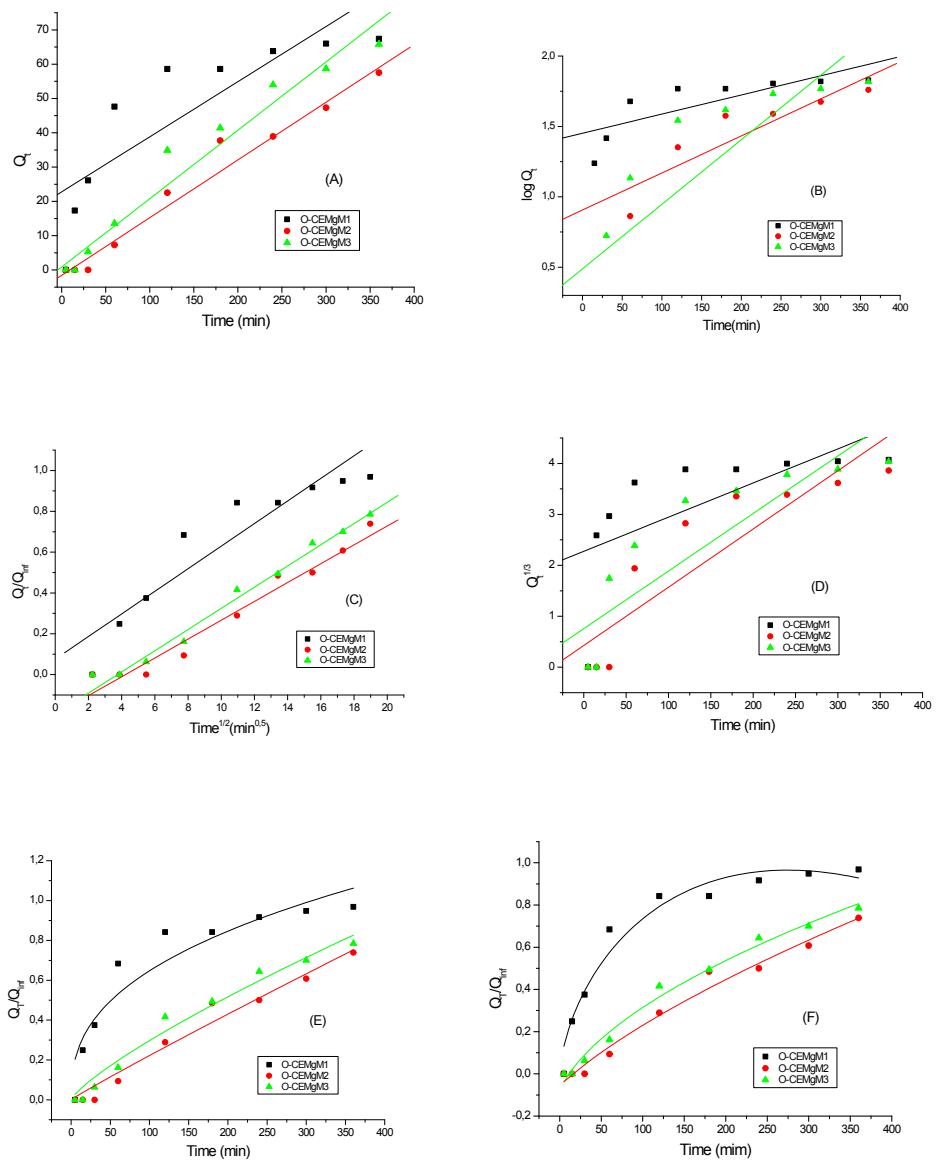


Figure S9: Mathematical models applied to maleimides release in phosphate buffer pH 7.4 to 37 °C of magnetic hydrogel. (A) Order zero, (B) first order (C) Higuchi, (D) Hixson-Crowell, (E)-Peppas and Korsmeyer (F) Peppas-Sahlin.

Table S10. Constant release and correlation coefficients of the mathematical equations applied in the release of the maleimides incorporated in the magnetic nanogel.

| | Zero order | | First order | | Higuchi | | Hixson-Crowell | | Korsmeyer-Peppas | | Peppas-Sahlin | | | | |
|--------|------------|--|-------------|--|---------|--|----------------|---|------------------|---------------------------|---------------|--------|--|--|--------|
| | r | K ₀ (min ⁻¹) | r | K ₁ (min ⁻¹) | r | K _H (min ^{-1/2}) | r | K _{HC} (min ^{-1/3}) | r | K (min ⁻¹) | n | r | K ₁ (min ⁻¹) | K ₂ (min ⁻¹) | m |
| pH 5.0 | | | | | | | | | | | | | | | |
| M1 | 0.9221 | 0.1616 | 0.8100 | 0.0022 | 0.9798 | 0.0603 | 0.7641 | 0.0075 | 0.9512 | 0.0491 | 0.5211 | 0.9900 | 0.0160 | -6.77x10 ⁻⁵ | 0.8305 |
| M2 | 0.9477 | 0.2579 | 0.8566 | 0.0022 | 0.9866 | 0.0740 | 0.8176 | 0.0112 | 0.9515 | 0.0254 | 0.6561 | 0.9909 | 0.0050 | -5.78x10 ⁻⁶ | 1.0557 |
| M3 | 0.9444 | 0.1667 | 0.8675 | 0.0024 | 0.9872 | 0.0553 | 0.7965 | 0.0078 | 0.9615 | 0.0304 | 0.5837 | 0.9913 | 0.0088 | -2.26x10 ⁻⁵ | 0.9082 |
| pH 7.4 | | | | | | | | | | | | | | | |
| M1 | 0.8296 | 0.1607 | 0.7785 | 0.0013 | 0.9223 | 0.05537 | 0.6612 | 0.0067 | 0.8969 | 0.1091 | 0.3867 | 0.9603 | 0.0465 | -5.6x10 ⁻⁴ | 0.6633 |
| M2 | 0.9936 | 0.1683 | 0.9127 | 0.0026 | 0.9838 | 0.04619 | 0.9048 | 0.0114 | 0.9718 | 0.0028 | 0.7518 | 0.9807 | 7.1x10 ⁻⁴ | -1.7x10 ⁻⁷ | 1.2609 |
| M3 | 0.9768 | 0.1995 | 0.7559 | 0.0046 | 0.9887 | 0.05196 | 0.8810 | 0.0113 | 0.9764 | 0.0075 | 0.7989 | 0.9909 | 0.0018 | -1.1x10 ⁻⁶ | 1.1335 |

Table S11: Release amount of maleimides (mg) from magnetic nanogel with and without application of magnetic fields, in phosphate buffer pH 5.0 and 37 °C after 360 min.

| | without EMF | with EMF |
|----------|-------------|----------|
| O-CEMgM1 | 1.76 | 2.55 |
| O-CEMgM2 | 1.87 | 2.70 |
| O-CEMgM3 | 2.61 | 2.79 |

Table S12: Constants release and correlation coefficients of the Korsmeyer-Peppas model applied in the release of the maleimides incorporated in the magnetic nanogel, in phosphate buffer pH 5.0, 37 °C.

| | r | K (min ⁻¹) | n |
|----------|--------|------------------------|--------|
| O-CEMgM1 | 0.9206 | 0.1274 | 0.3725 |
| O-CEMgM2 | 0.9260 | 0.0453 | 0.5633 |
| O-CEMgM3 | 0.9789 | 0.0791 | 0.4578 |