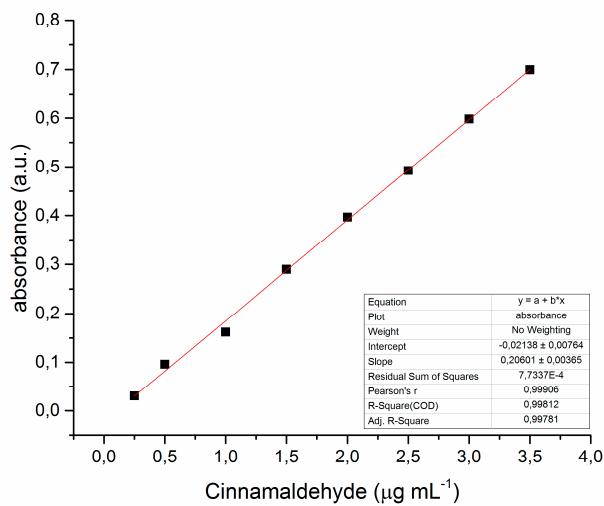
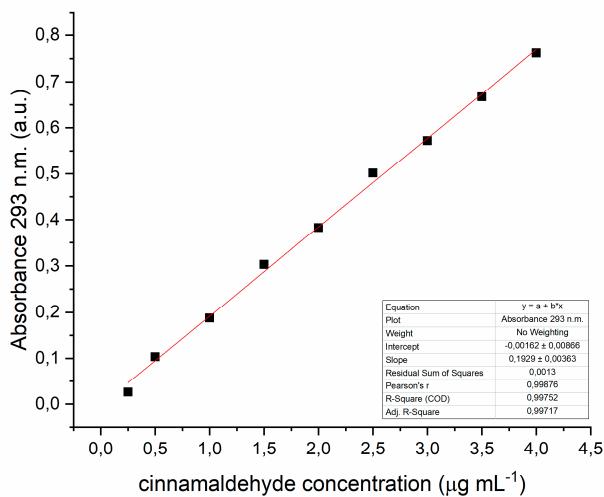


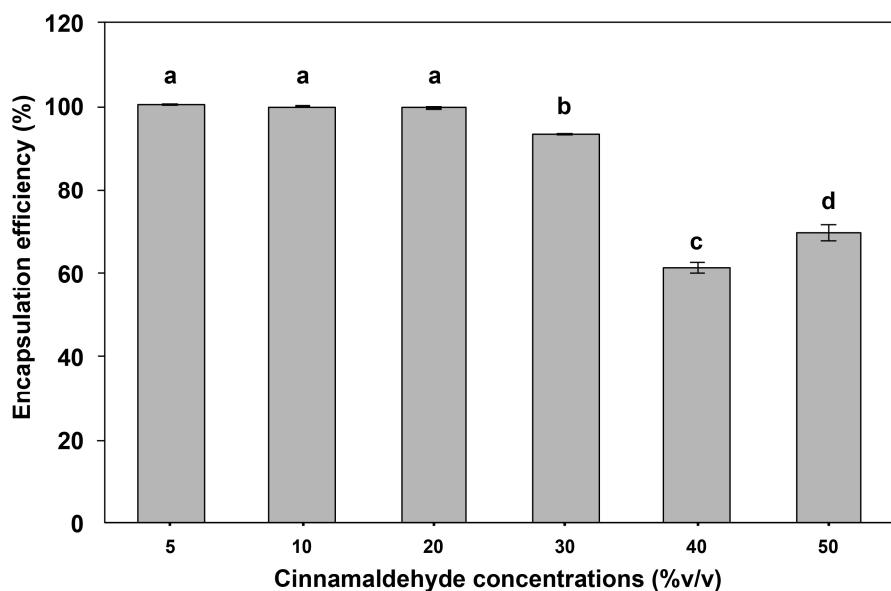
## Supplementary material S1



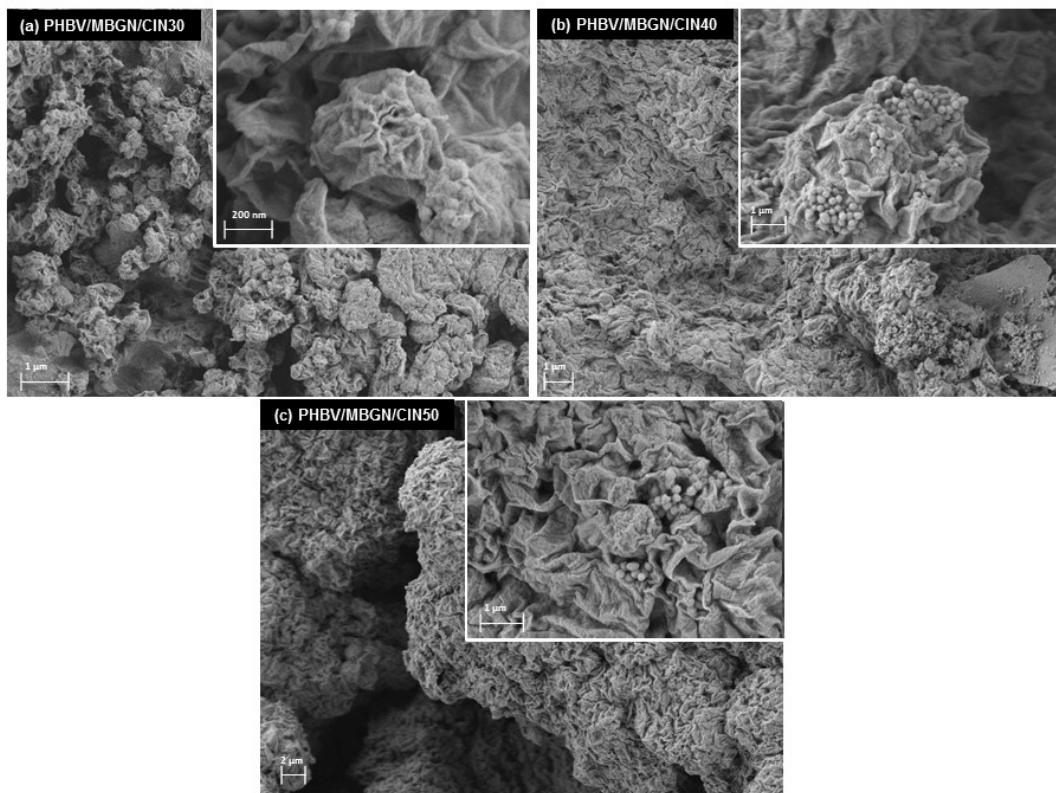
**Figure S1.** Linear regression equation, corresponding to the calibration curve of cinnamaldehyde in methanol by means of the absorbance UV-vis spectrophotometry at 287 nm, ranging from 0.25  $\mu\text{g mL}^{-1}$  to 3.5  $\mu\text{g mL}^{-1}$



**Figure S2.** Linear regression equation, corresponding to the calibration curve of cinnamaldehyde in simulated body fluid (SBF) by means of the absorbance UV-vis spectrophotometry at 293 nm, ranging from 0.25  $\mu\text{g mL}^{-1}$  to 3.5  $\mu\text{g mL}^{-1}$



**Figure S3.** Encapsulation efficiency of PHBV/MBGN microspheres as function of CIN concentration ranging from 5 to 50% (v/v). Experimental data are reported as mean  $\pm$  standard deviation.  $n = 3$ . Means followed by the different letters within columns indicate a significant difference at  $p < 0.05$  using Duncan's new multiple range test.



**Figure S4.** Scanning electron micrographs of (a) PHBV/MBGN/CIN30, (b) PHBV/MBGN/CIN40, and (c) PHBV/MBGN/CIN50 microspheres.

**Table S1.** the raw data of diameter, polydispersity index and zeta potential of blank and cinnamaldehyde-loaded microspheres.

sample	Size ( $\mu\text{m}$ )					PDI					Zeta potential (mV)				
	#1	#2	#3	AVG	STV	#1	#2	#3	AVG	STV	#1	#2	#3	AVG	STV
PHBV/MBGN	6.8	5.1	6.5	6.1	0.7	1.0	0.7	0.9	0.9	0.1	-20.2	-21.2	-20.8	-20.7	0.4
PHBV/MBGN/CIN5	5.9	6.2	9.3	7.2	1.5	0.5	0.4	0.4	0.4	0.1	-20.7	-21.4	-21.8	-21.3	0.5
PHBV/MBGN/CIN10	10.4	10.0	13.6	11.4	1.6	0.6	0.3	0.8	0.6	0.2	-20.1	-21.1	-20.0	-20.4	0.5
PHBV/MBGN/CIN20	13.7	14.5	9.2	12.5	2.3	0.6	0.8	0.2	0.5	0.2	-15.9	-10.0	-10.6	-12.2	2.7

**Table S2.** the raw data of encapsulation efficiency of PHBV/MBGN microspheres on cinnamaldehyde concentration ranging from 5 % (v/v) to 50% (v/v)

Cinnamaldehyde concentration (% v/v)	Encapsulation efficiency (%)				
	#1	#2	#3	AVG	STV
5%	99.9635	99.9563	99.9579	99.9592 <sup>a</sup>	0.0031
10%	99.8503	99.8481	99.7997	99.8327 <sup>a</sup>	0.0233
20%	99.2044	99.3072	99.2678	99.2598 <sup>a</sup>	0.0423
30%	55.8538	48.3190	49.0014	51.0581 <sup>b</sup>	3.4025
40%	60.5063	62.8804	60.1217	61.1695 <sup>c</sup>	1.2199
50%	66.4520	70.4902	71.6485	69.5303 <sup>d</sup>	2.2274

**Table S3.** the raw data of *in vitro* cinnamaldehyde cumulative release (%) of PHBV/MBGN/CIN5, PHBV/MBGN/CIN10, and PHBV/MBGN/CIN20 microspheres in phosphate buffer solution within 336 h

The different time point (h)	sample	Cinnamaldehyde cumulative release ( $\mu\text{g mL}^{-1}$ )				
		#1	#2	#3	AVG	STV
0.5	PHBV/MBGN/CIN5	0.20	0.43	0.40	0.35	0.10
0.5	PHBV/MBGN/CIN10	0.79	0.77	0.56	0.71	0.11
0.5	PHBV/MBGN/CIN20	9.12	10.04	8.39	9.19	0.68
1	PHBV/MBGN/CIN5	0.87	1.29	1.18	1.11	0.18
1	PHBV/MBGN/CIN10	1.90	1.99	1.78	1.89	0.09
1	PHBV/MBGN/CIN20	11.97	12.91	11.04	11.97	0.76
2	PHBV/MBGN/CIN5	1.32	1.99	1.70	1.67	0.27
2	PHBV/MBGN/CIN10	2.65	2.78	2.58	2.67	0.08
2	PHBV/MBGN/CIN20	14.41	15.86	13.23	14.50	1.08
4	PHBV/MBGN/CIN5	1.54	2.36	1.90	1.94	0.33
4	PHBV/MBGN/CIN10	3.06	3.23	2.93	3.07	0.12
4	PHBV/MBGN/CIN20	15.97	17.70	14.95	16.21	1.14
6	PHBV/MBGN/CIN5	1.54	2.56	1.90	2.00	0.42
6	PHBV/MBGN/CIN10	3.21	3.48	3.05	3.25	0.18
6	PHBV/MBGN/CIN20	16.61	18.71	15.53	16.95	1.32
8	PHBV/MBGN/CIN5	1.54	2.73	1.90	2.06	0.50
8	PHBV/MBGN/CIN10	3.21	3.71	3.05	3.32	0.28
8	PHBV/MBGN/CIN20	16.94	19.37	15.83	17.38	1.48
24	PHBV/MBGN/CIN5	1.66	2.98	2.03	2.23	0.55
24	PHBV/MBGN/CIN10	3.43	4.02	3.24	3.57	0.33
24	PHBV/MBGN/CIN20	17.45	20.16	16.36	17.99	1.60
48	PHBV/MBGN/CIN5	1.66	3.19	2.03	2.30	0.65
48	PHBV/MBGN/CIN10	3.60	4.28	3.43	3.77	0.37
48	PHBV/MBGN/CIN20	17.76	20.69	16.66	18.37	1.70

72	PHBV/MBGN/CIN5	1.66	3.34	2.03	2.34	0.72
72	PHBV/MBGN/CIN10	3.60	4.47	3.43	3.83	0.46
72	PHBV/MBGN/CIN20	17.98	21.07	16.83	18.62	1.79
168	PHBV/MBGN/CIN5	1.66	3.54	2.03	2.41	0.81
168	PHBV/MBGN/CIN10	3.60	4.72	3.56	3.96	0.54
168	PHBV/MBGN/CIN20	18.18	21.49	17.05	18.91	1.88
336	PHBV/MBGN/CIN5	1.79	3.72	2.03	2.52	0.86
336	PHBV/MBGN/CIN10	3.72	4.94	3.68	4.11	0.59
336	PHBV/MBGN/CIN20	18.33	21.79	17.22	19.11	1.95

**Table S4.** the raw data of Dissolution Data Modeling and goodness of fit of zero order model

Parameters	#1	#2	#3
N_observed	11	11	11
DF	10	10	10
R_obs-pre	0.5184	0.5426	0.5116
Rsqr	-5.2249	-5.6337	-18.2879
Rsqr_adj	-5.2249	-5.6337	-18.2879
MSE	4157.6866	4093.0410	5355.4208
MSE_root	64.4801	63.9769	73.1807
Weighting	1	1	1
SS	41576.8665	40930.4099	53554.2084
WSS	41576.8665	40930.4099	53554.2084
AIC	118.9883	118.8159	121.7729
MSC	-2.0104	-2.0740	-3.1413

**Table S5.** the raw data of Dissolution Data Modeling and goodness of fit of first order model

Parameters	#1	#2	#3
N_observed	11	11	11
DF	10	10	10
R_obs-pre	0.9655	0.9645	0.9617
Rsqr	0.8680	0.8432	0.7773
Rsqr_adj	0.8680	0.8432	0.7773
MSE	88.1826	96.7320	61.8218
MSE_root	9.3906	9.8352	7.8627
Weighting	1	1	1
SS	881.8260	967.3205	618.2179
WSS	881.8260	967.3205	618.2179
AIC	76.6019	77.6198	72.6953
MSC	1.8429	1.6711	1.3203

**Table S6.** the raw data of Dissolution Data Modeling and goodness of fit of Higuchi model

Parameters	#1	#2	#3
N_observed	11	11	11
DF	10	10	10
R_obs-pre	0.6667	0.6929	0.6707
Rsqr	-2.2395	-2.4213	-9.9452
Rsqr_adj	-2.2395	-2.4213	-9.9452
MSE	2163.7177	2110.9723	3039.0105

MSE_root	46.5158	45.9453	55.1272
Weighting	1	1	1
SS	21637.1769	21109.7227	30390.1052
WSS	21637.1769	21109.7227	30390.1052
AIC	111.8039	111.5324	115.5406
MSC	-1.3572	-1.4118	-2.5747

**Table S7.** the raw data of Dissolution Data Modeling and goodness of fit of Hixson model

Parameters	#1	#2	#3
N_observed	11	11	11
DF	10	10	10
R_obs-pre	0.6665	0.6955	0.6737
Rsqr	-3.0093	-3.2426	-12.4093
Rsqr_adj	-3.0093	-3.2426	-12.4093
MSE	2677.8719	26177.7142	3723.1951
MSE_root	51.7482	51.1636	61.0180
Weighting	1	1	1
SS	26778.7195	26177.1422	37231.9512
WSS	26778.7195	26177.1422	37231.9512
AIC	114.1490	113.8991	117.7741
MSC	-1.5704	-1.6270	-2.7778

**Table S8.** the raw data of Dissolution Data Modeling and goodness of fit of Korsmeyer-Peppas model

Parameters	#1	#2	#3
N_observed	11	11	11
DF	9	9	9
R_obs-pre	0.8271	0.8497	0.8605
Rsqr	0.6812	0.7192	0.7394
Rsqr_adj	0.6457	0.6880	0.7105
MSE	236.6112	192.5241	80.3846
MSE_root	15.3822	13.8753	8.9657
Weighting	1	1	1
SS	2129.5004	1732.7166	723.4613
WSS	2129.5004	1732.7166	723.4613
AIC	88.3001	86.0319	76.4245
MSC	0.7795	0.9064	0.9813

**Table S9.** the raw data of antibacterial activity of PHBV/MBGN/CIN5, PHBV/MBGN/CIN10, and PHBV/MBGN/CIN20 microspheres on *S. aureus* and *E. coli*

The different time point (h)	sample	Relative bacterial viability (%)				
		#1	#2	#3	AVG	STV
<i>S. aureus</i>						
3	PHBV/MBGN	99.10	99.40	98.60	99.03	0.40
3	PHBV/MBGN/CIN5	83.20	89.60	92.90	88.57	4.93
3	PHBV/MBGN/CIN10	79.00	80.90	83.90	81.27	2.47
3	PHBV/MBGN/CIN20	50.90	55.80	46.20	50.97	4.80
6	PHBV/MBGN	98.30	97.20	97.40	97.63	0.59
6	PHBV/MBGN/CIN5	84.60	91.70	80.10	85.47	5.85
6	PHBV/MBGN/CIN10	77.40	82.30	83.70	81.13	3.31
6	PHBV/MBGN/CIN20	13.40	24.90	28.00	22.10	7.69
24	PHBV/MBGN	98.60	94.90	96.10	96.53	1.89
24	PHBV/MBGN/CIN5	56.00	58.90	64.50	59.80	4.32
24	PHBV/MBGN/CIN10	57.40	58.10	56.90	57.47	0.60
24	PHBV/MBGN/CIN20	20.80	17.80	14.20	17.60	3.30
<i>E. coli</i>						
3	PHBV/MBGN	99.20	97.10	98.30	98.20	1.05
3	PHBV/MBGN/CIN5	88.70	87.10	78.90	84.90	5.26
3	PHBV/MBGN/CIN10	85.20	81.60	84.40	83.73	1.89
3	PHBV/MBGN/CIN20	19.80	17.70	13.60	17.03	3.15
6	PHBV/MBGN	98.30	98.10	97.50	97.97	0.42
6	PHBV/MBGN/CIN5	81.90	79.90	83.30	81.70	1.71
6	PHBV/MBGN/CIN10	81.20	77.80	83.30	80.77	2.78
6	PHBV/MBGN/CIN20	19.90	17.70	15.30	17.63	2.30
24	PHBV/MBGN	98.30	97.70	95.90	97.30	1.25
24	PHBV/MBGN/CIN5	62.60	65.20	61.10	62.97	2.07
24	PHBV/MBGN/CIN10	56.20	62.10	60.20	59.50	3.01
24	PHBV/MBGN/CIN20	45.80	43.50	41.70	43.67	2.06

**Table S10.** the raw data of cytotoxicity test of PHBV/MBGN/CIN5, PHBV/MBGN/CIN10, and PHBV/MBGN/CIN20 microspheres on osteosarcoma MG-63 cells

The different dose ( $\mu\text{g mL}^{-1}$ )	sample	Relative cell viability (%)				
		#1	#2	#3	AVG	STV
Day 1						
1000	PHBV/MBGN/CIN5	87.41	90.23	91.35	89.66	1.66
1000	PHBV/MBGN/CIN10	85.15	87.41	84.59	85.71	1.22
1000	PHBV/MBGN/CIN20	72.91	76.35	70.94	73.40	2.24
100	PHBV/MBGN/CIN5	98.68	93.05	96.99	96.24	2.36
100	PHBV/MBGN/CIN10	87.97	89.10	89.66	88.91	0.70
100	PHBV/MBGN/CIN20	83.74	79.31	77.83	80.30	2.51
10	PHBV/MBGN/CIN5	100.38	99.25	101.50	100.38	0.92
10	PHBV/MBGN/CIN10	93.61	96.43	93.61	94.55	1.33
10	PHBV/MBGN/CIN20	85.22	87.68	95.07	89.33	4.19
1	PHBV/MBGN/CIN5	98.12	94.74	93.05	95.30	2.11
1	PHBV/MBGN/CIN10	92.48	93.05	94.17	93.23	0.70

<b>1</b>	PHBV/MBGN/CIN20	87.19	67.49	118.23	<b>90.97</b>	20.89
<b>Day 5</b>						
<b>1000</b>	PHBV/MBGN/CIN5	70.77	64.20	66.70	67.22	2.71
<b>1000</b>	PHBV/MBGN/CIN10	56.37	65.45	57.62	59.81	4.02
<b>1000</b>	PHBV/MBGN/CIN20	50.51	56.63	53.95	53.70	2.51
<b>100</b>	PHBV/MBGN/CIN5	88.94	91.44	92.07	90.81	1.35
<b>100</b>	PHBV/MBGN/CIN10	87.37	91.41	84.24	87.68	2.95
<b>100</b>	PHBV/MBGN/CIN20	94.13	92.98	89.16	92.09	2.13
<b>10</b>	PHBV/MBGN/CIN5	104.28	101.77	98.96	101.67	2.17
<b>10</b>	PHBV/MBGN/CIN10	100.84	98.02	96.14	98.33	1.93
<b>10</b>	PHBV/MBGN/CIN20	99.49	99.87	104.08	101.15	2.08
<b>1</b>	PHBV/MBGN/CIN5	103.03	100.21	103.34	102.19	1.41
<b>1</b>	PHBV/MBGN/CIN10	99.90	98.96	100.21	99.69	0.53
<b>1</b>	PHBV/MBGN/CIN20	104.85	105.99	110.59	107.14	2.48