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Title: Novel antimicrobial superporous cross-linked chitosan/pyromellitimide benzoyl thiourea hydrogels

Author: Nadia A. Mohamed Nahed A Abd. El-Ghany Mona

M. Fahmy

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## ACCEPTED MANUSCRIPT

| Novel antimicrobial superporous cross-linked chitosan/pyromellitimide benzoyl                     | 1  |
|---|----|
| thiourea hydrogels  | 2  |
| Nadia A. Mohamed*, Nahed A. Abd El-Ghany, Mona M. Fahmy   | 3  |
| Department of Chemistry, Faculty of Science, Cairo University, Giza, 12613, Egypt;                | 4  |
| E-Mails: <u>dr_nahed_055@yahoo.com</u> (N.A.A.); <u>m.fahmy17@yahoo.com</u> (M.M.F.)              | 5  |
| *Author to whom correspondence should be addressed; E-Mail: namadm@hotmail.com                    | 6  |
| (N.A.M.)  | 7  |
| Abstract  | 8  |
| In this work, chitosan (CS) was cross-linked with different amounts of pyromellitimide            | 9  |
| benzoyl thiourea moieties. The structure of the cross-linked CS was confirmed by                  | 10 |
| elemental analyses, FTIR and <sup>1</sup> H- NMR spectroscopy. The cross-linking process proceeds | 11 |
| via reacting of the amino groups of CS with the isothiocyanate groups of the N,N'-bis [4-         | 12 |
| (isothiocyanate carbonyl)phenyl] pyromellitimide cross-linker. The amount of the cross-           | 13 |
| linker was varied with respect to CS to produce four new pyromellitimide benzoyl                  | 14 |
| thiourea cross-linked CS (PIBTU-CS) hydrogels designated as PIBTU-CS-1, PIBTU-CS-                 | 15 |
| 2, PIBTU-CS-3, and PIBTU-CS-4 of increasing cross-linking degree percent of 11, 22, 44            | 16 |
| and 88%, respectively. The scanning electron microscopy observation indicates the                 | 17 |
| extremely porous structure of the hydrogels. XRD results showed that the crystallinity of         | 18 |
| CS was decreased upon cross-linking. The four hydrogels exhibit a higher antibacterial            | 19 |
| activity on Bacillus subtilis and Streptococcus pneumoniae as Gram positive bacteria and          | 20 |
| against Escherichia coli as Gram negative bacteria and higher antifungal activity on              | 21 |
| Aspergillus fumigatus, Syncephalastrum racemosum and Geotricum candidum than that of              | 22 |
| the parent CS as shown from their higher inhibition zone diameters and their lower MIC            | 23 |
| values. The swell ability of the hydrogel as well as their antimicrobial activity increased       | 24 |
| with increasing cross-linking density.  | 25 |
| Keywords: Chitosan hydrogels, Synthesis, Characterization, Swell ability,                         | 26 |
| Antimicrobial activity  | 27 |
|   | 28 |

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