

The Modification of Poly amidoamine (PAMAM-G_{0.5}) by Cytosine

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Received 2012

ABSTRACT

The half generation Poly(amidoamine) (PAMAM) dendrimers was synthesized and characterized and modification by heterocyclic DNA base (Cytosine). The conjugated Cytosine modified detected by FT-IR, ¹H NMR, ¹³C NMR and Mass spectroscopy analysis.

Keywords: Component; Polyamidoamine; PAMAM; Cytosine

1. Introduction

Dendrimers are molecules with three-dimensional structure of certain branches around a central core which were synthesized by processing stage or repeated synthesis. Also dendrimers have been identified alike the arborols, starburst, ascade and Cauliflower polymers. Dendrimers are Unique molecular structure of polymer materials like balls that contain a Initiator core, repeat units and Surface terminal group. Those are spherical configuration in high Generation. For the first time we have invented new models for determination molecular weight dendrimers AB₂-type [9,12]. Dendrimer polyamidoamine (PAMAM) as one of commercial ones [10,13] due to the controllable mass, the water solubility, and the possibility of surface functionality, has been widely used in the biomedical and genetics [10]. The large numbers of surface functional groups on dendrimer's outer shell can be modified or conjugated with a variety of interesting guest molecules [11,14]. PAMAM was modified by different molecules, in this study PAMAM was modified to cytosine. The modification PAMAM molecule properties is changed proportion to PAMAM molecule, in hence modification PAMAM can be conjugated with targeting molecules, dyes and drugs for different usage in science [5-8].

2. Experimental Process

2.1. Synthesis PAMAM Dendrimers Half Generation (G_{0.5}), EDA Core (PAMAM(G_{0.5})-EDA(Core), Z₀=2)

The Synthesis PAMAM dendrimers half generation (G_{0.5}), EDA Core (PAMAM(G_{0.5})-EDA(Core), Z₀=2) A solution of freshly distilled 1,2-diaminoethane (5 g, 5.5 ml, 0.083 mol) in methanol (20 ml) was added dropwise to a stirred solution of methylacrylate (35 g, 37 ml, 0.407 mol) in methanol (20 ml), under nitrogen, over a period of 2 h. The final mixture was stirred for 30 min at 0°C and then allowed to warm to room temperature and stirred for a further 24 h. The solvent was removed under reduced pressure at 40° C using a rotary evapo-

rator and the resulting yellow oil viscose under vacuum (10-1 mm Hg, 50°C) overnight to give PAMAM(G_{0.5})-EDA(Core) (a), the final product (3.64 g, 91.5%). The synthesis (a) in **Figure 1** is shown. 1738 cm⁻¹ is ester C=O stretching absorption. ¹³C-NMR ester groups carbon has appeared in 173.4 ppm. ¹H-NMR ester methyl group protons was appeared in 2.51 ppm. Full spectral data and characterizations (a) is given in **Table 1**.

2.2. Conjugated of Cytosine with G_{0.5} PAMAM Dendrimer

Cytosine (4.5 mmol) was dissolved in H₂O at PAMAM(G_{0.5})-EDA(Core)(a) (4.95 mmol) was dissolved in MeOH, Then was added drop wise to the cytosine solution, at a molar ratio of cytosine/(a) 4:1. After complete addition the mixture was stirred for 48h at room temperature and the resultant light yellow solid vacuum dried (10-1 mm Hg, 50°C) overnight to give the cytosine with PAMAM-EDA(Core) (b) (71.9%). It was detectable by FT-IR, ¹H NMR, ¹³C NMR, and Mass spectroscopy analysis. The synthesis (b) in **Figure 2** is shown.

3. Results and Discussion

FT-IR spectra (a), (b) and cytosine was shown in **Figure 3 i, ii, iii** respectively. ester groups absorption in 1735.8 cm⁻¹ which not presented in the cytosine spectra (**Figure 4 iii**) whereas disappeared thoroughly in (b) (**Figure 4 ii**). at CH₂ groups anti-symmetrical stretching was obtained (a) at 2954 cm⁻¹. of -NH and stretching vibrations was shown at 3285 cm⁻¹ (**Figure 4i**). -NH stretching vibrations amide was shown at 1648

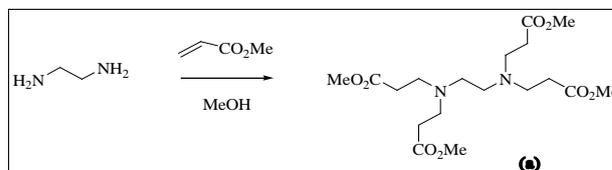


Figure 1. Synthesis PAMAM (G_{0.5}).

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Table 1. Spectral mode assignment for a, b and Cytosine.

Spectral data	PAMAM G _{0.5}	Cytosine	PAMAM G _{0.5} (Cyt) ₄		
	C=O stretching vibration ester groups	1735	C=O Stretching vibration amide 1725	C=O Stretching vibration amide 1566	
FT-IR (cm ⁻¹)	C-O stretching ester groups	1203	C=N stretching vibration 1666	N-H bending vibration amide 1234.4	
	C-N stretching vibration amine groups	1041	C-N stretching vibration 1234	C-N stretching vibration 1651	
¹³ CNMR (ppm)	C=O	173.1	C=O	166.9	C=O (core) 176.9
	C-C=O		C-NH ₂		C-NH-C=O 164.7
	CH ₃	32.6 51.9	C-NH-C=O	171 158.1	
¹ HNMR (ppm)	CH ₂ -C=O	2.46	CH (ring)	5.62	CH ₂ -C=O 2.28
	CH ₃ (end groups)	3.64			NH-C=O (new bond) 7.3
Mass (m/z)	404	111			483-564-644-720

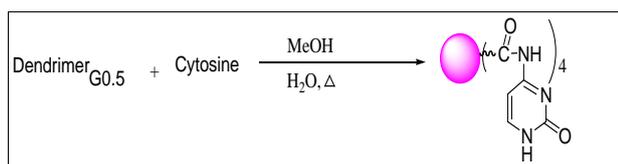


Figure 2. Synthesis PAMAM-EDA(Core)-Cytosine (b).

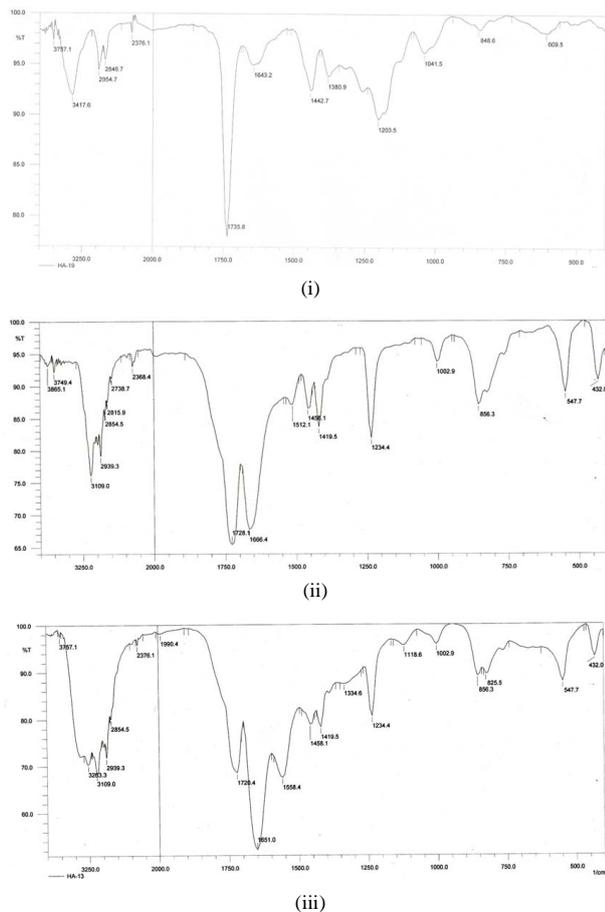


Figure 4. FT-IR spectral (a), (b) and Cytosine in this figure; i, ii, iii respectively.

cm-1 and 1558 cm-1 for amide I & II respectively. Also N-H bending vibrations amide was shown at 1648 and 1558 cm-1. Spectral data and characterizations (a), (b) and cytosine are given in Table 1.

4. Conclusions

In summary, conjugating cytosine with half generation of PAMAM dendrimer was successfully synthesized and characterized. PAMAMG_{0.5}-Cytosine Systematic mass spectral analysis has shown that nearly theoretical masses are obtained for product (b) (or the mass spectra of this compound displayed molecular ion peaks at the appropriate m/z values or mass spectra on the other hand, give precise molecular weight information to compare with mathematically predicted molecular masses.), mass spectrometry of that gave a mass of 720 m/z (theoretical mass: 724m/z). Overall the spectral data clearly indicate the desired cytosine with PAMAM-EDA(Core) (b) is formed.

REFERENCES

- [1] S.M. Buck, Y.E.L. Koo, E. Park, H. Xu, M.A. Philbert, M.A. Brasuel; R. Kopelman,. Optochemical nanosensor PEBBLES: photonic explorers for bioanalysis with biologically localized embedding, Current Opinion in Chemical Biology 2004, 8, 540-546.
- [2] A.J. Haes, R.P. Van Duyne, Preliminary studies and potential applications of localized surface plasmon resonance spectroscopy in medical diagnostics, Expert Review of Molecular Diagnostics 2004, 4, 527-537.
- [3] J.L. West, N.J. Halas, Engineered nanomaterials for biophotonics applications: improving sensing, imaging, and therapeutics, Annual Review of Biomedical Engineering 2003, 5, 285-292.
- [4] J.M.J. Fréchet, D.A. Tomalia Dendrimers and Other Dendritic Polymers, John Wiley & Sons, West Sussex 2001
- [5] K.K. Ong, A.L. Jenkins; R. Chen, D.A. Tomalia, H.D. Durst, Dendrimer enhanced immunosensors for biological detection, Analytica Chimica Acta 2001, 444, 143-148.

- [6] C. Dufes, W.N. Keith, A. Bilsland, I. Proutski; J.F. Uchegbu; Schatzlein, A.G. Synthetic anticancer gene medicine exploits intrinsic antitumor activity of cationic vector to cure established tumors, *Cancer Research* 2005, 65, 8079–8084.
- [7] A. D'Emanuele, Attwood, D. Dendrimer-drug interactions, *Advanced Drug Delivery Reviews* 2005, 57, 147–2162.
- [8] Venditto, V.J.; Regino, C.A.S.; Brechbiel, M.W. PAMAM dendrimer based macromolecules as improved contrast agents, *Molecular Pharmaceutics* 2005, 2, 302–311.
- [9] A. H. Massoudi , H. Vahedi, O. Louie , S.Sajjadifar ; *E – Journal of Chemistry* 2009, Vol. 6(3)681-684.
- [10] D.A. Tomalia, *Chem. Today* 2005, 23,41.
- [11] W. Pei, Zh. Xin-Han, W. Zhi-Yu, M. Min, Li. Xu, N. Qian, Generation 4 polyamidoamine dendrimers is a novel candidate of nano-carrier for gene delivery agents in breast cancer treatment, *Cancer Letters*. 2010, 298, 34–49.
- [12] O.Louie, A.H. Massoudi, H. Vahedi, S.Sajjadifar, "Determination of molecular weight and molecular radius of the polyamido carboxylic acid dendrimer using generation numbers", *Polymer*, 2009, 50, 5605–5607
- [13] O. Louie et al, PAMAM Megamer (G2-G2) as a versatile in gene delivery , *Clinical Biochemistry*, *Clinical Biochemistry*, Volume 44, Issue 13, Supplement, September 2011, Page S281-S282
- [14] A.M. Massoudi, H. Vahedi, O. Louie, S. Sajjadifar, S. Damavandi, *Der Chemica Sinica*, 2011, 2 (4):312-315