

Synthesis of Lakshminine and Antiproliferative Testing of Related Oxoisoaporphines

Vicente Castro-Castillo,^{*,†,‡} Marco Rebollo-Fuentes,[‡] Cristina Theoduloz,[▽] and Bruce K. Cassels,^{‡,§}

Faculty of Basic Sciences, Metropolitan Educational Sciences University, Av. J.P. Alessandri 774, Ñuñoa, Santiago, Chile; Department of Chemistry, Faculty of Sciences, University of Chile, Santiago, Chile; Faculty of Health Sciences, University of Talca, Talca, Chile; Millennium Institute for Cell Dynamics and Biotechnology, Santiago, Chile

* To whom correspondence should be addressed: tel.: +56 2 241 2494; fax: +56 2 239 3932; e-mail: vicente.castro@umce.cl.

† Faculty of Basic Sciences.

‡ Faculty of Sciences.

▽ Faculty of Health Sciences.

§ Millennium Institute.

Supporting information

Contents

T1. In vitro Antiproliferative Activity of Amino-1-azabenzanthrones **1** and **2**, the Related Compounds **6**, **8**, **10**, and **11**, and Etoposide (**ETO**), Towards Normal and Cancerous Human Cell Lines.

S1. ^1H -NMR (400 MHz, CDCl_3) spectrum of 2,3-dihydro-5-methoxy-6-nitro- $7H$ -dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-6-nitrobenzanthrone, **7**).

S2. ^{13}C -NMR (100 MHz, $\text{DMSO}-d_6$) spectrum of 2,3-dihydro-5-methoxy-6-nitro- $7H$ -dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-6-nitrobenzanthrone, **7**).

S3. ^1H -NMR (400 MHz, CDCl_3) spectrum of 2,3-dihydro-5-methoxy-4-nitro- $7H$ -dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-4-nitrobenzanthrone, **9**).

S4. ^{13}C -NMR (100 MHz, $\text{DMSO}-d_6$) spectrum of 2,3-dihydro-5-methoxy-4-nitro- $7H$ -dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-4-nitrobenzanthrone, **9**).

S5. ^1H -NMR (400 MHz, CDCl_3) spectrum of 5-methoxy-6-nitro- $7H$ -dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-6-nitrobenzanthrone, **8**).

S6. ^{13}C -NMR (100 MHz, $\text{DMSO}-d_6$) spectrum of 5-methoxy-6-nitro- $7H$ -dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-6-nitrobenzanthrone, **8**).

S7. ^1H -NMR (400 MHz, CDCl_3) spectrum of 6-amino-5-methoxy- $7H$ -dibenzo[*de,h*]quinolin-7-one (lakshminine, **1**).

S8. ^{13}C -NMR (100 MHz, CDCl_3) spectrum of 6-amino-5-methoxy- $7H$ -dibenzo[*de,h*]quinolin-7-one (lakshminine, **1**).

S9. ^1H -NMR (400 MHz, $\text{DMSO}-d_6$) spectrum of 4-amino-5-methoxy- $7H$ -dibenzo[*de,h*]quinolin-7-one (**2**).

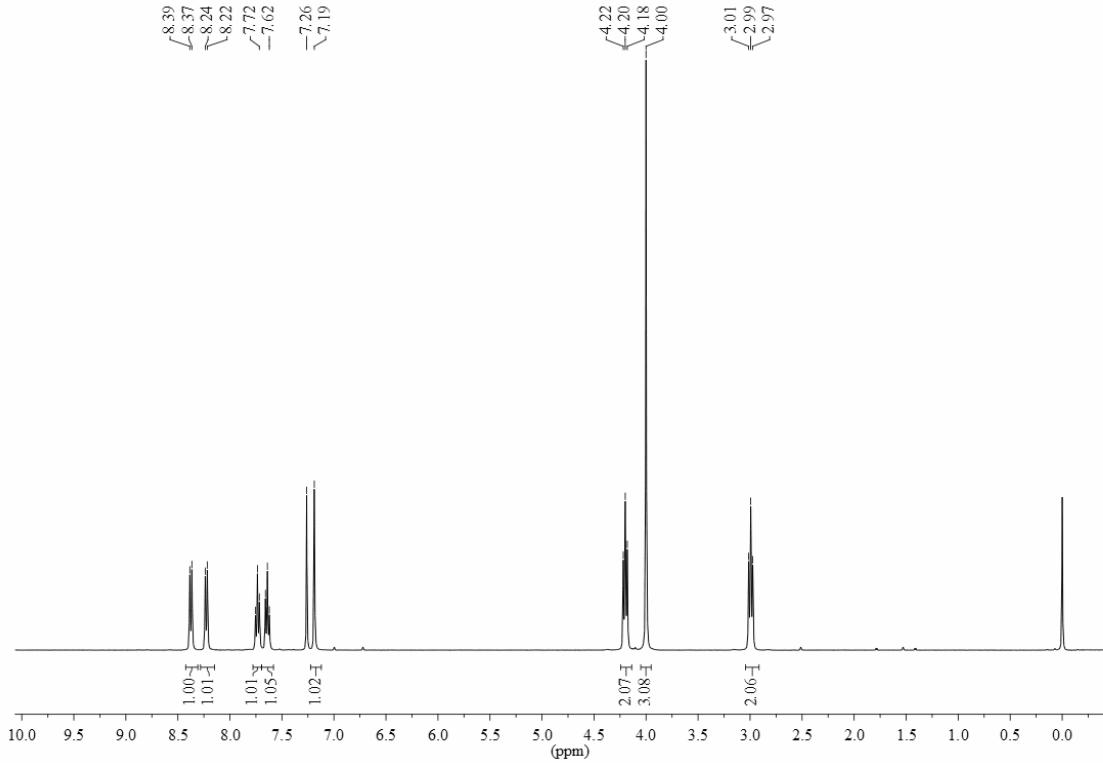
S10. ^{13}C -NMR (100 MHz, $\text{DMSO}-d_6$) spectrum of 4-amino-5-methoxy- $7H$ -dibenzo[*de,h*]quinolin-7-one (**2**).

Compound	Cell line IC ₅₀ ± SEM (μM)			
	MRC-5 ^a	AGS ^b	SK-MES-1 ^c	J82 ^d
1	17.9 ± 0.7	78.6 ± 3.9	43.3 ± 3.4	37.6 ± 2.6
2	37.0 ± 1.9	> 100	> 100	> 100
6	> 100	> 100	> 100	> 100
8	80.0 ± 4.8	47.7 ± 2.9	> 100	48.3 ± 2.4
10	10.2 ± 0.5	4.5 ± 0.3	17.1 ± 1.0	25.4 ± 1.3
11	41.6 ± 2.9	32.9 ± 1.6	> 100	93.6 ± 5.7
ETO	3.9 ± 0.2	0.36 ± 0.02	2.5 ± 0.2	2.8 ± 0.2

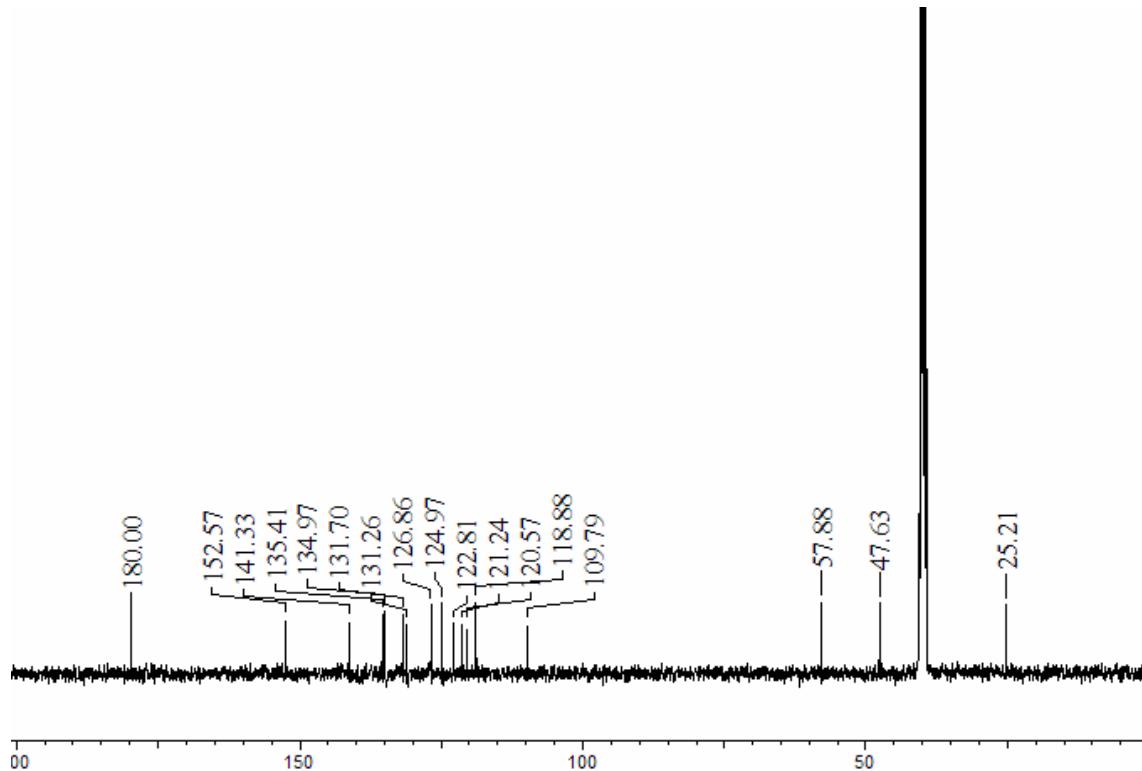
Results are shown as the means of the concentrations (μM) inhibiting cell growth by 50% ± SEM (three independent determinations). All compounds were tested as free bases.

^a MRC-5: normal lung fibroblasts (CCL-171); ^bAGS: gastric adenocarcinoma cells (CRL-1739); ^cSK-MES-1: lung cancer cells (HTB-58); ^dJ82: bladder carcinoma (HTB-1).

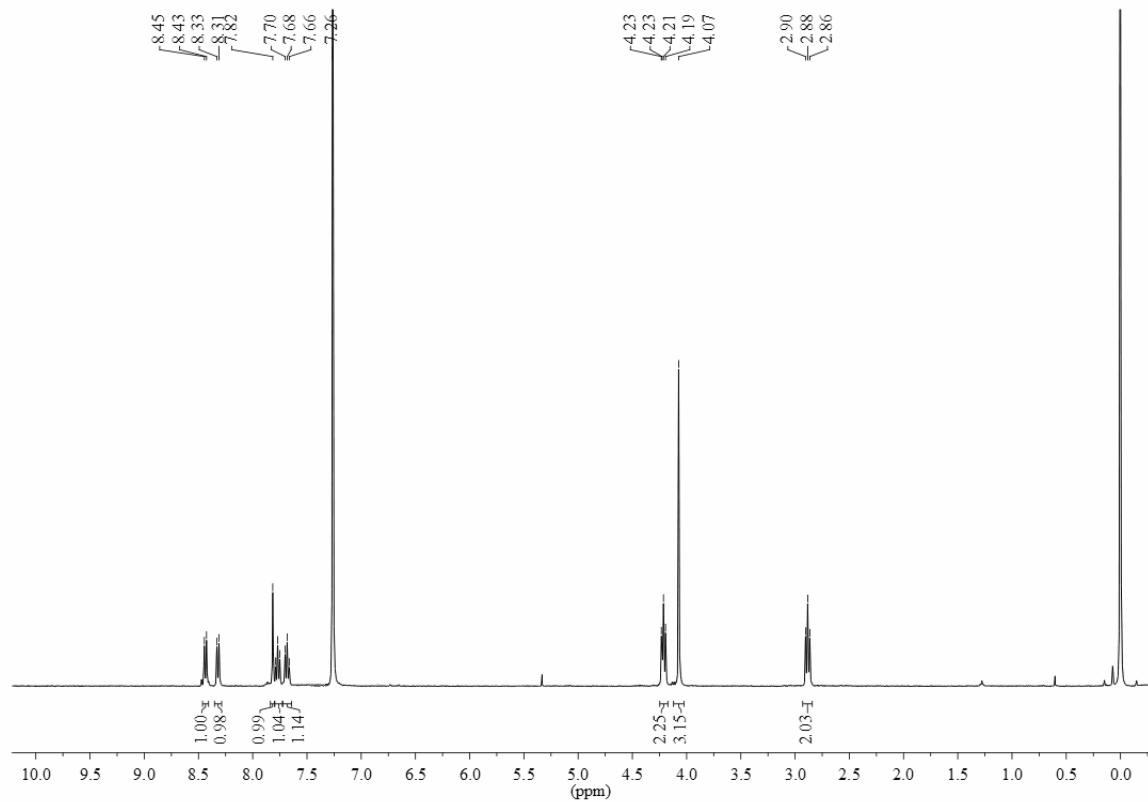
Table 1. In vitro Antiproliferative Activity of Amino-1-azabenzanthrones **1** and **2**, the Related Compounds **6**, **8**, **10**, and **11**, and Etoposide (**ETO**), Towards Normal and Cancerous Human Cell Lines.



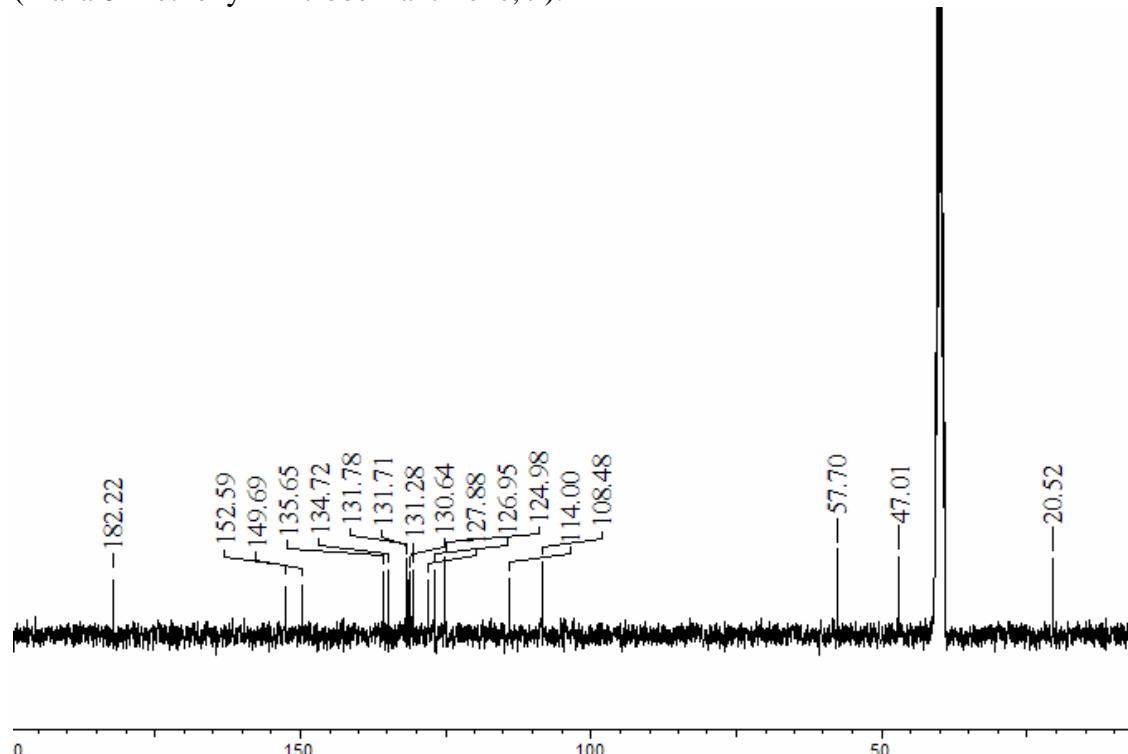
S1. ^1H -NMR spectrum of 2,3-dihydro-5-methoxy-6-nitro-7*H*-dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-6-nitrobenzanthrone, 7).



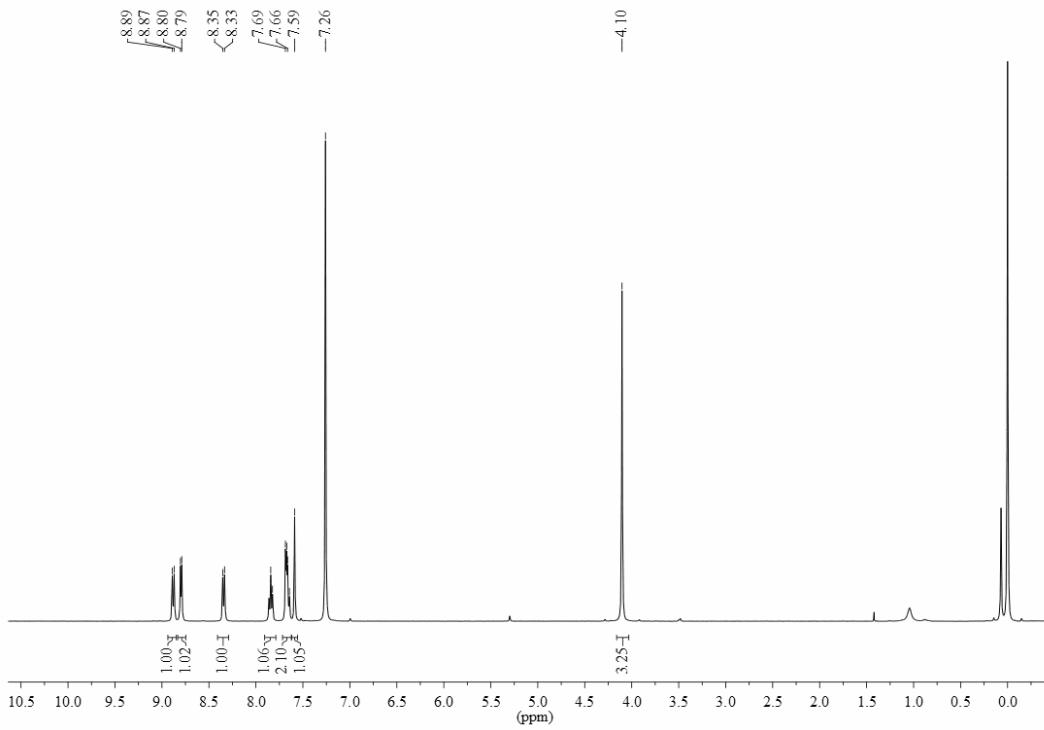
S2. ^{13}C -NMR spectrum of 2,3-dihydro-5-methoxy-6-nitro-7*H*-dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-6-nitrobenzanthrone, 7).



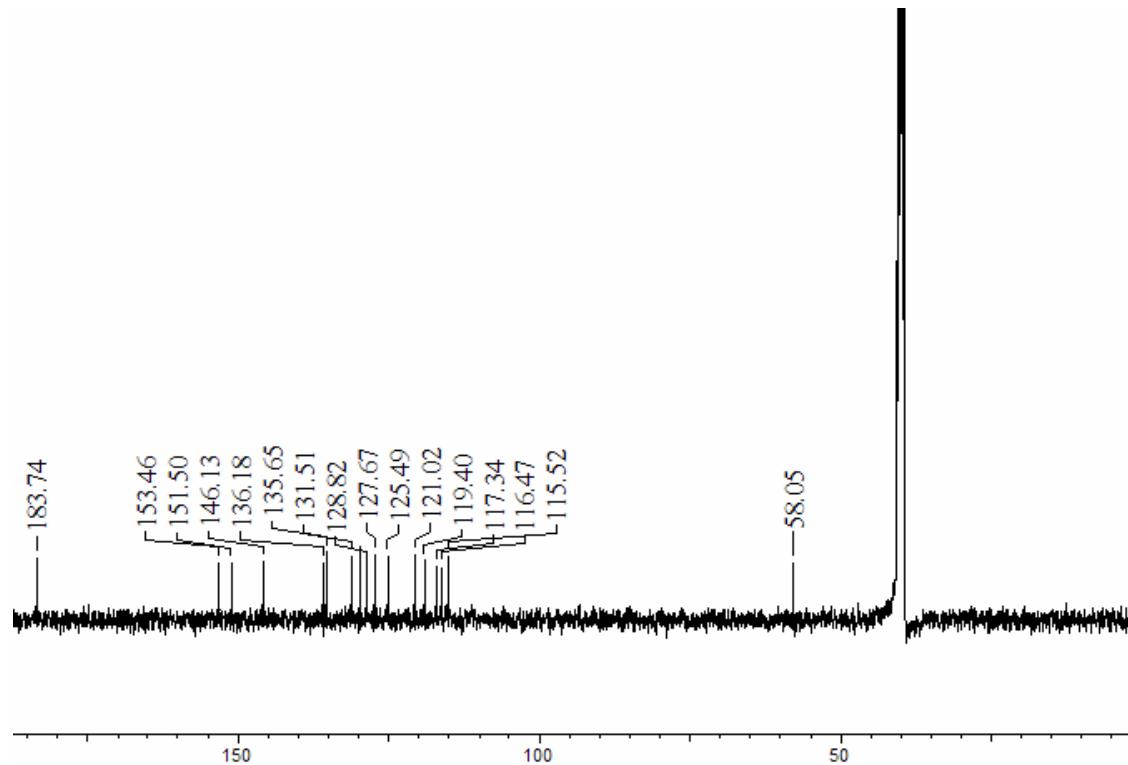
S3. ^1H -NMR spectrum of 2,3-dihydro-5-methoxy-4-nitro-7*H*-dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-4-nitrobenzanthrone, **9**).



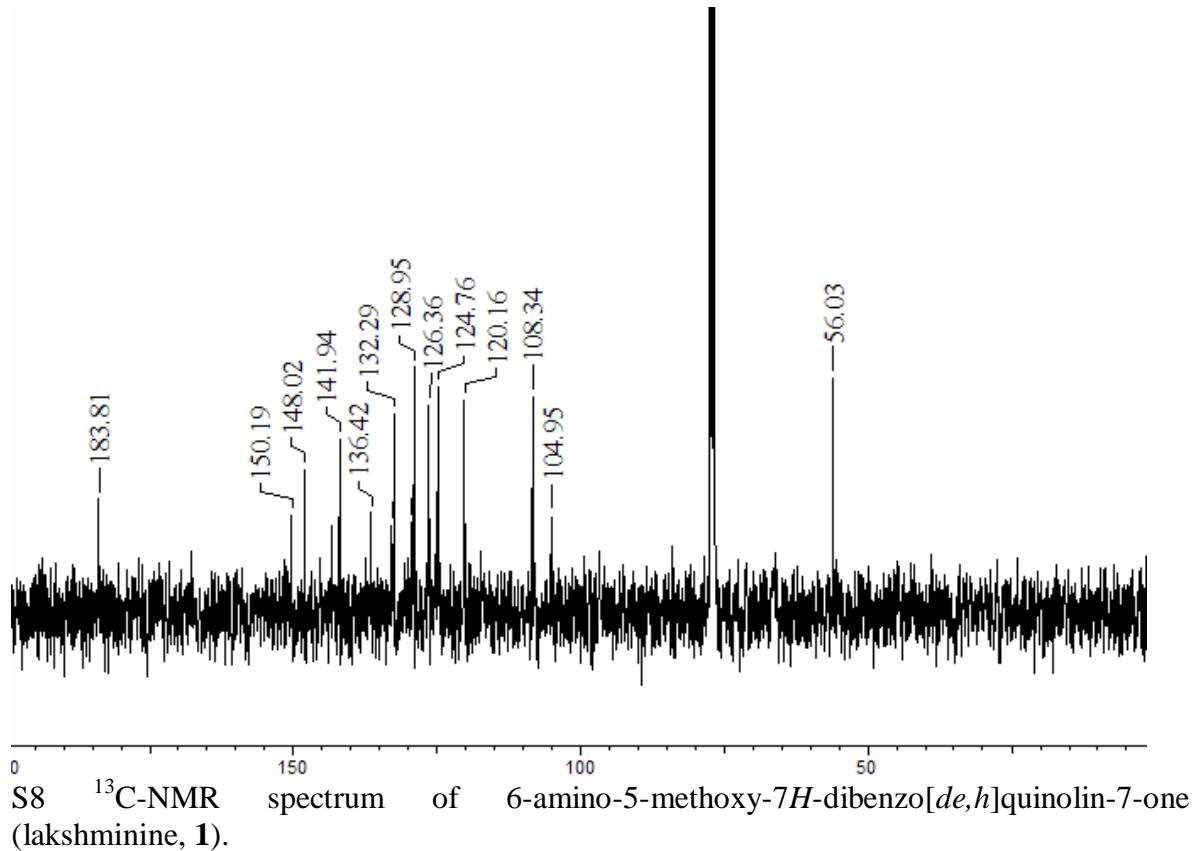
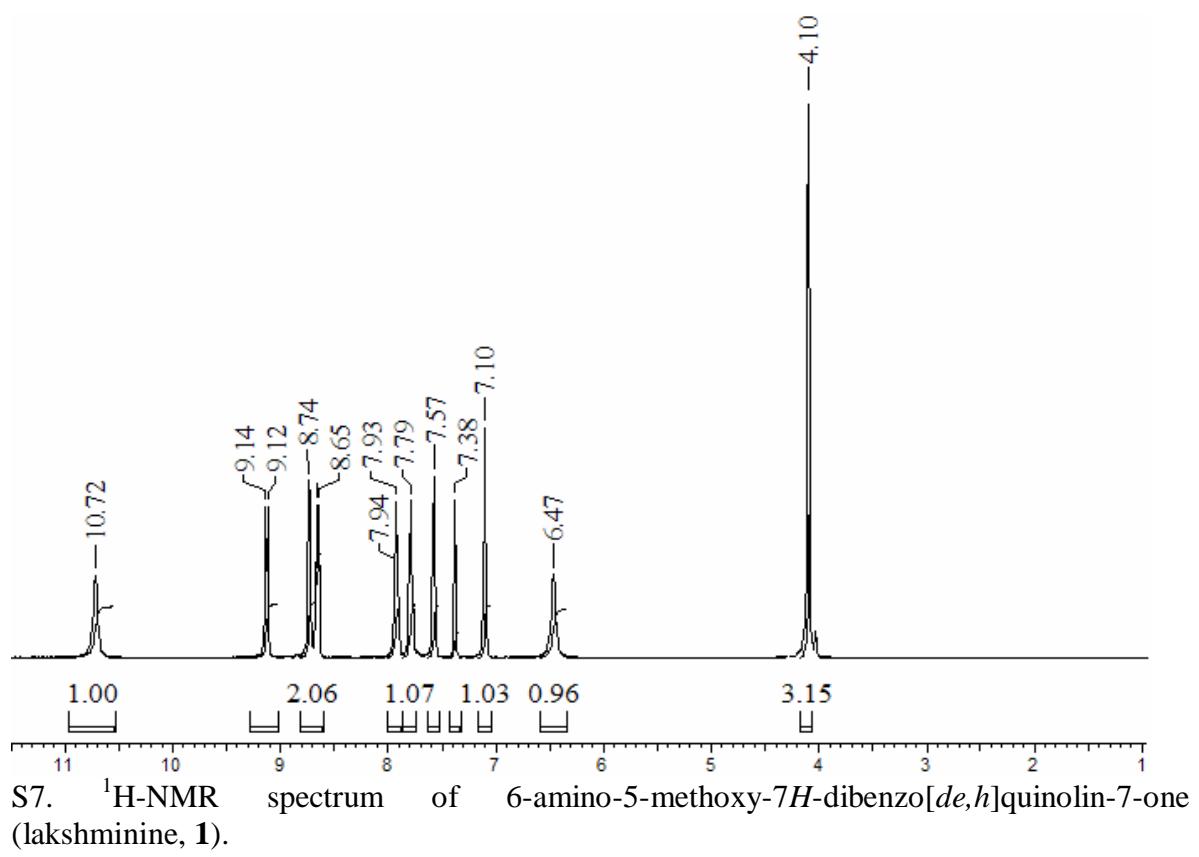
S4. ^{13}C -NMR spectrum of 2,3-dihydro-5-methoxy-4-nitro-7*H*-dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-4-nitrobenzanthrone, **9**).

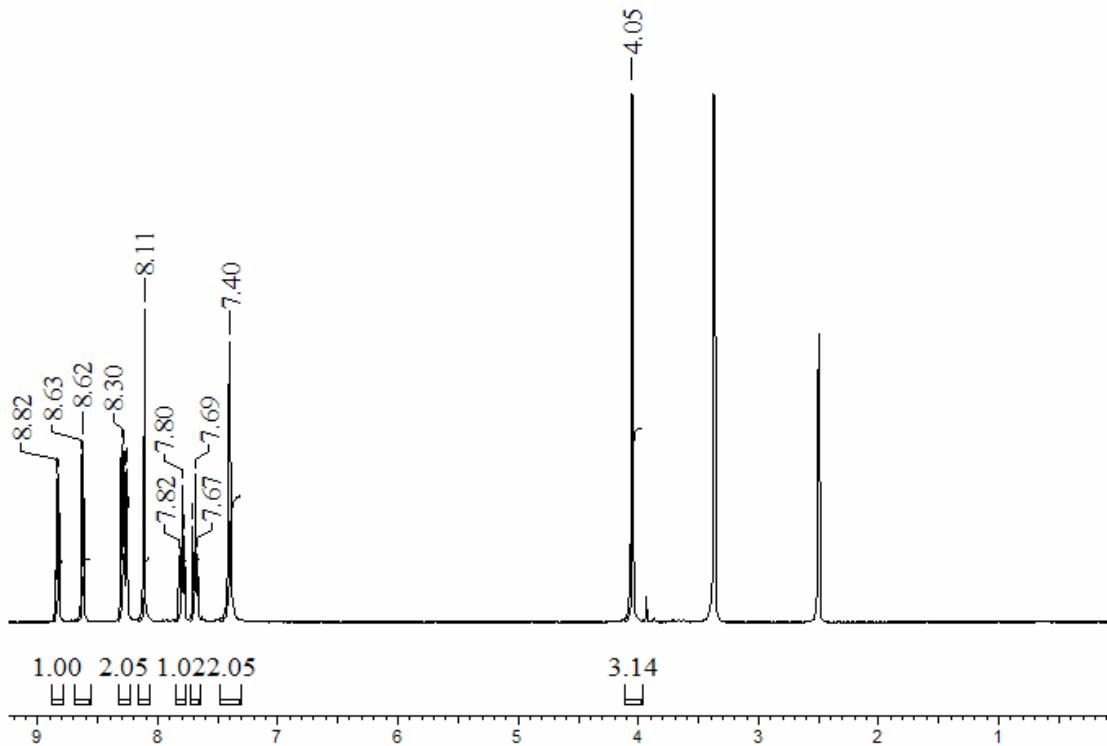


S5. ¹H-NMR spectrum of 5-methoxy-6-nitro-7*H*-dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-6-nitrobenzanthrone, **8**).

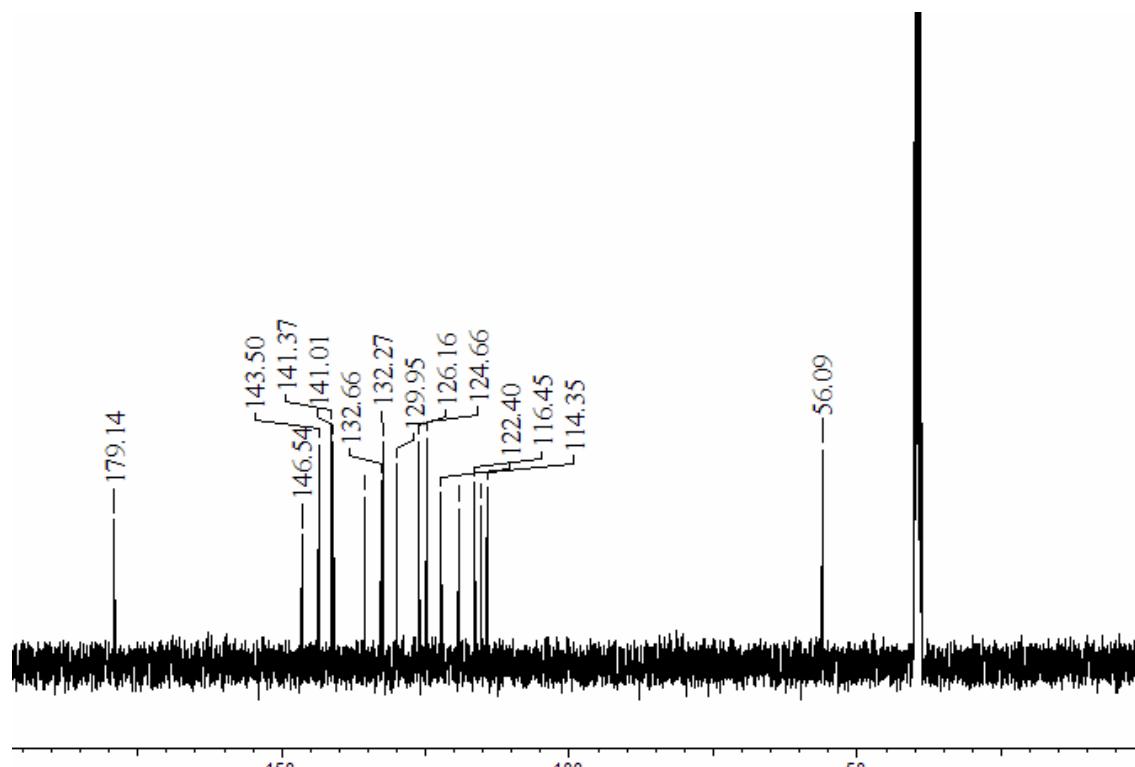


S6. ¹³C-NMR spectrum of 5-methoxy-6-nitro-7*H*-dibenzo[*de,h*]quinolin-7-one (1-aza-5-methoxy-6-nitrobenzanthrone, **8**).





S9. ¹H-NMR spectrum of 4-amino-5-methoxy-7H-dibenzo[*de,h*]quinolin-7-one (**2**).



S10. ¹³C-NMR spectrum of 4-amino-5-methoxy-7H-dibenzo[*de,h*]quinolin-7-one (**2**).