

Supplementary Materials: Backstabbing P-gp: Side-Chain Cleaved Ecdysteroid 2,3-Dioxolanes Hyper-Sensitize MDR Cancer Cells to Doxorubicin without Efflux Inhibition

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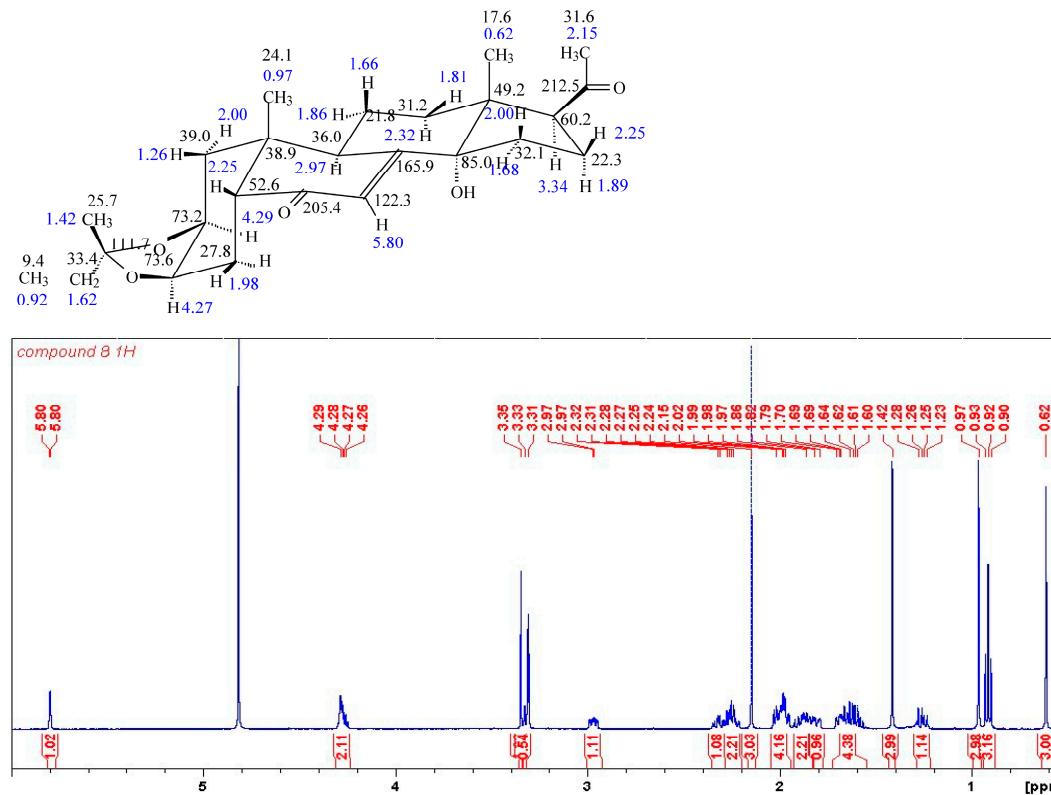


Figure S1. ^1H -NMR spectrum of compound 8.

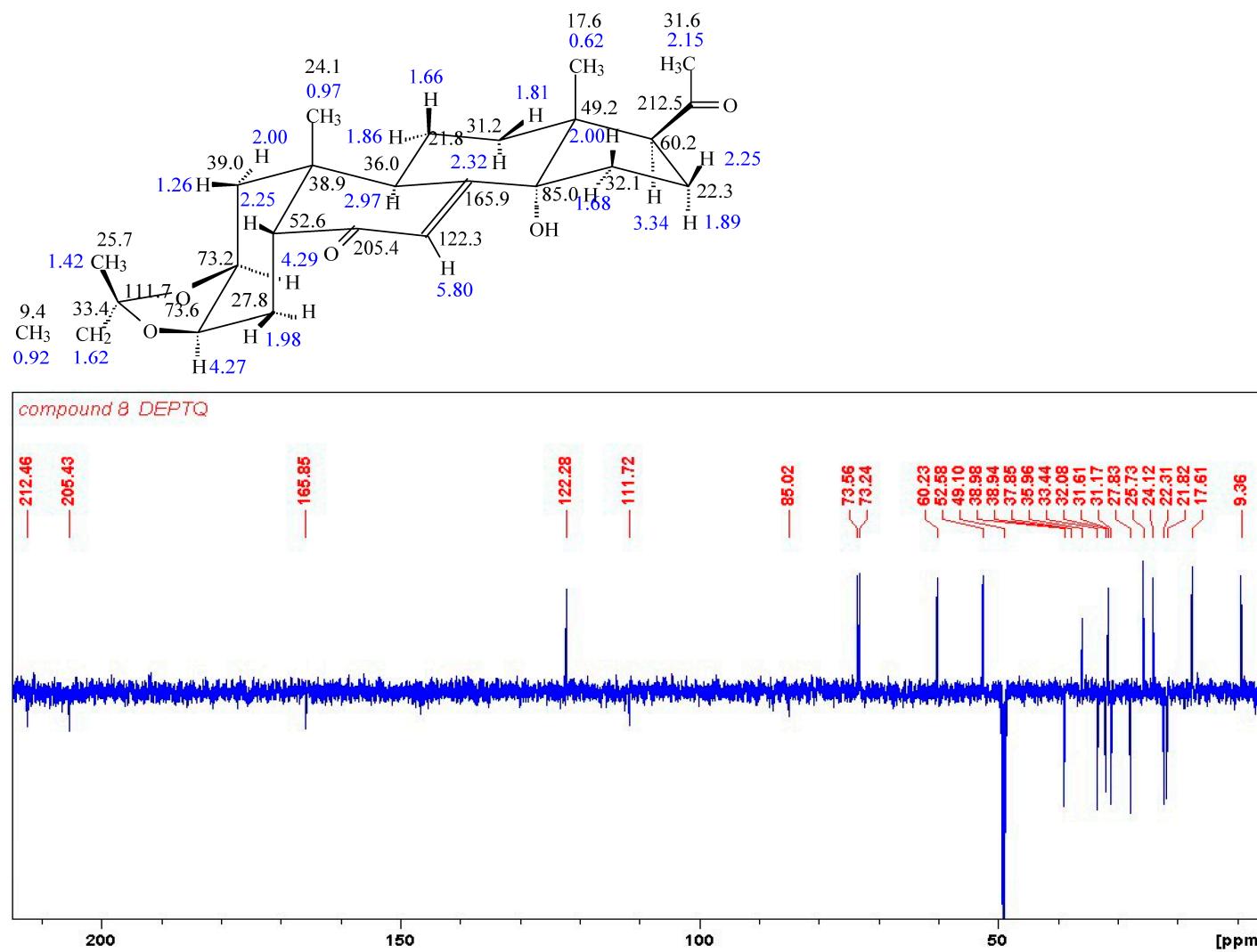


Figure S2. DEPTQ spectrum of compound 8.

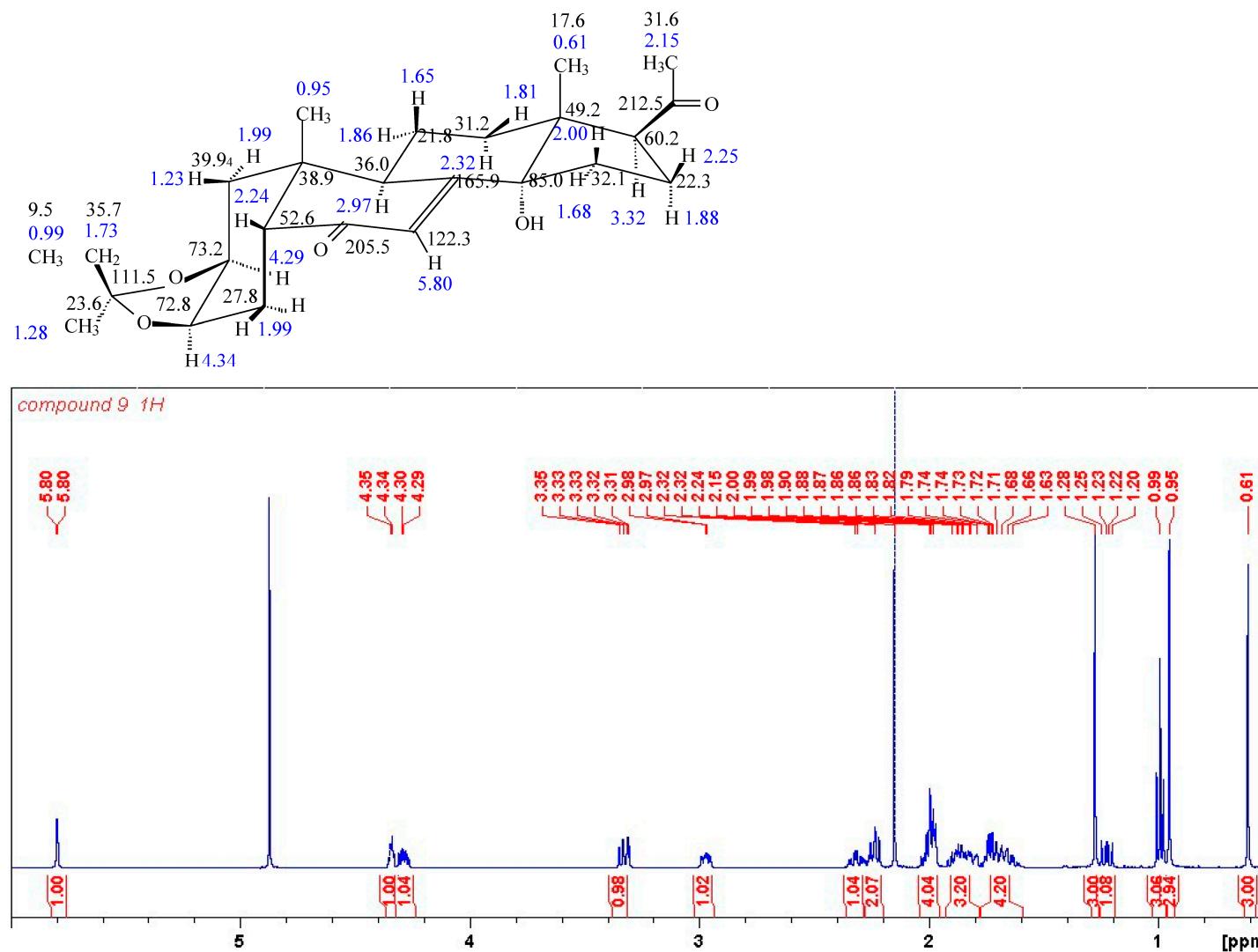


Figure S3. ^1H -NMR spectrum of compound 9.

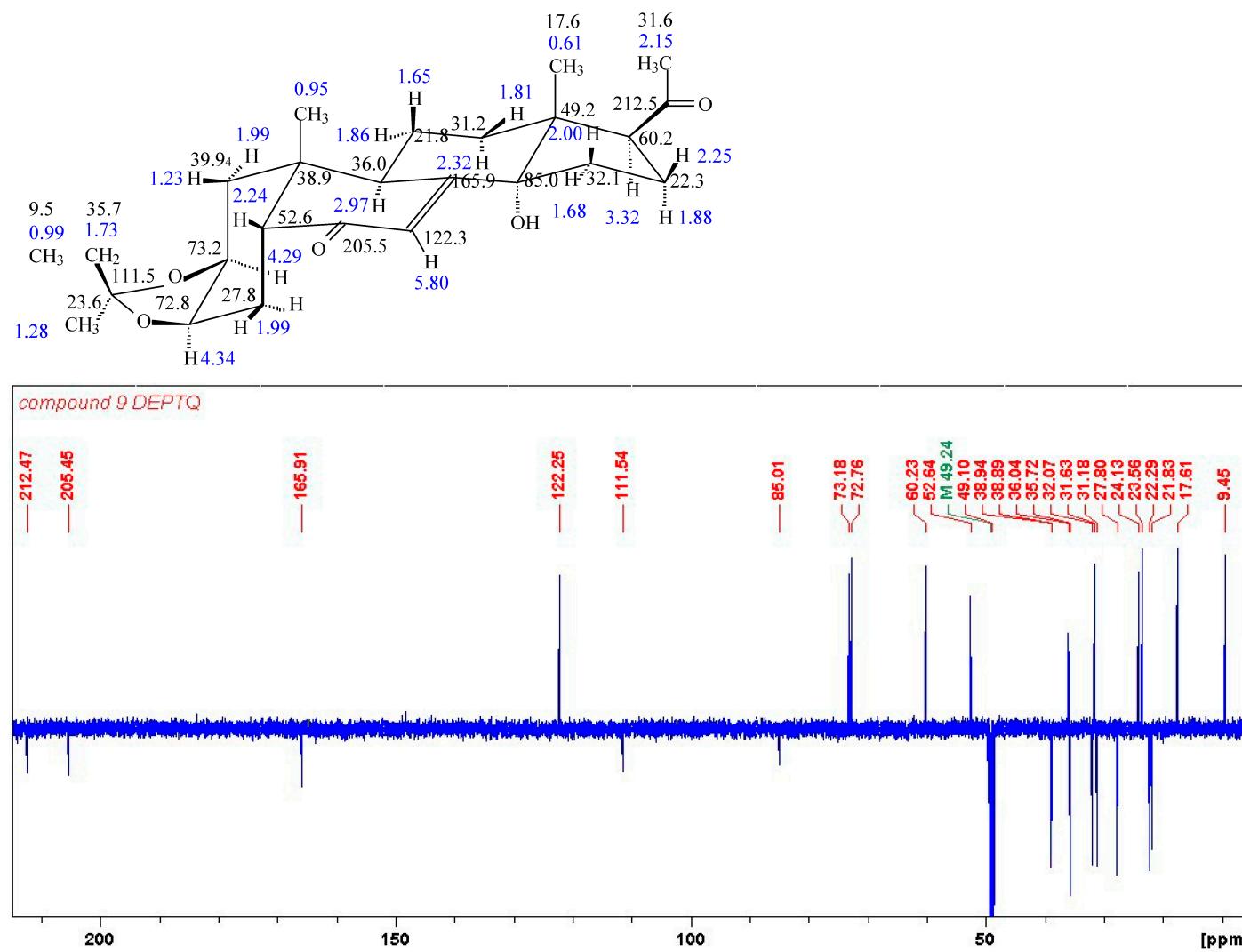


Figure S4. DEPTQ spectrum of compound 9.

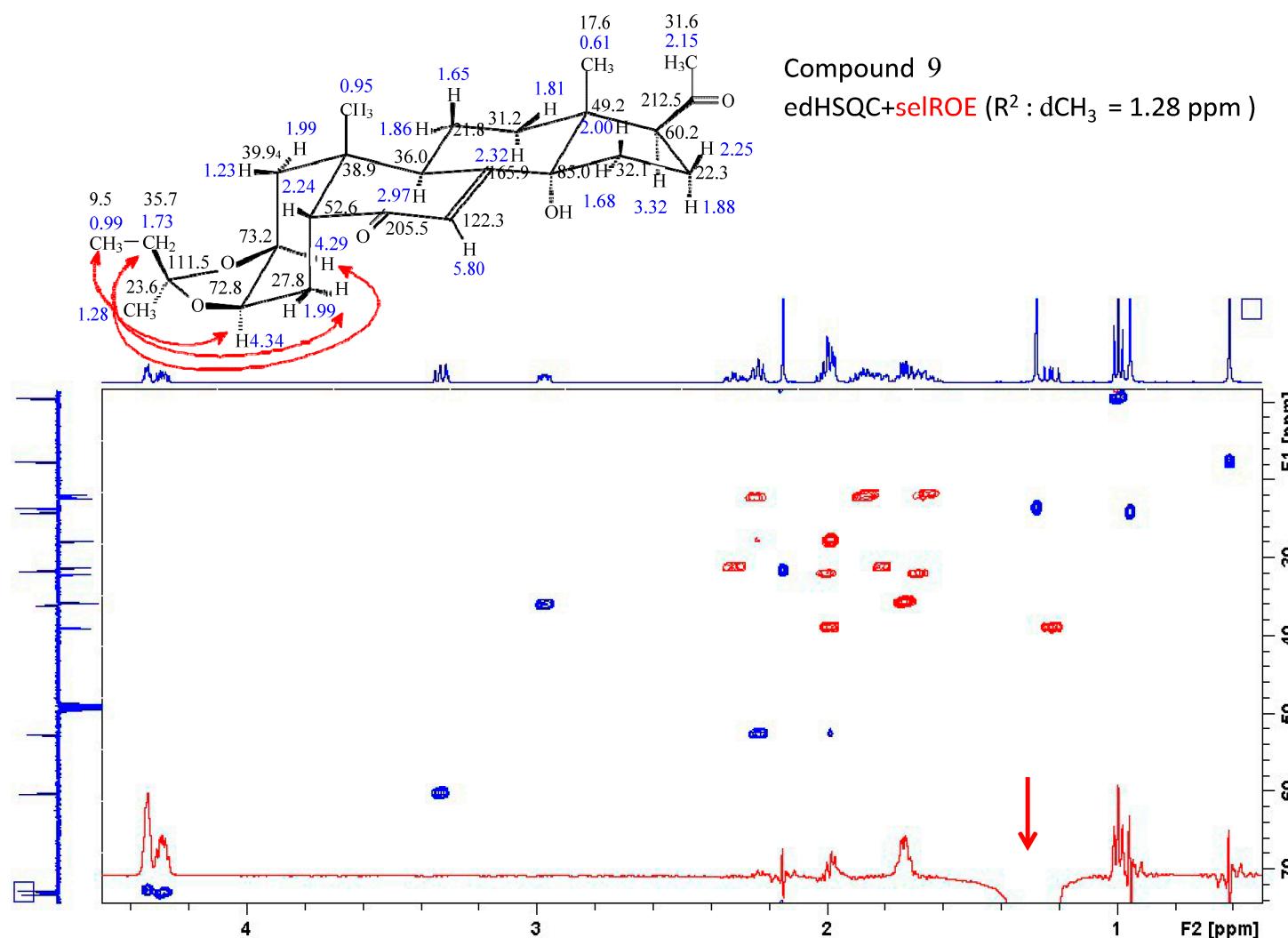


Figure S5. Confirmation of the stereochemistry of compound **9** at C-22.

Table S1. Cytotoxicity of doxorubicin alone and in combination with 10 or 25 μ M of compounds **1–10** on the L5178 and L5178_{MDR} cell lines.

Cell Line	Doxorubicin Alone	Ecdysteroid	IC ₅₀ of Doxorubicin (μ M)										
			20DA	1	2	3	4	5	6	7	8	9	10
L5178	0.41 ± 0.02	10 μ M	0.27 ± 0.03 **	0.37 ± 0.02	0.34 ± 0.03	0.33 ± 0.02	0.26 ± 0.02 **	0.28 ± 0.02 **	0.21 ± 0.003 **	0.18 ± 0.005 **	0.19 ± 0.02 **	0.34 ± 0.04	0.24 ± 0.03 **
		25 μ M	0.17 ± 0.01 **	0.29 ± 0.01	*0.23 ± 0.02 **	0.15 ± 0.006 **	0.17 ± 0.01 **	0.20 ± 0.02 **	0.15 ± 0.02 **	0.12 ± 0.003 **	0.11 ± 0.003 **	0.26 ± 0.07	*0.23 ± 0.03 **
L5178 _{MDR}	11.8 ± 0.64	10 μ M	1.17 ± 0.08 **	13.22 ± 0.44	3.80 ± 0.10 **	2.74 ± 0.66 **	1.74 ± 0.11 **	6.81 ± 0.33 **	1.56 ± 0.14 **	0.43 ± 0.02 **	2.60 ± 0.11 **	2.23 ± 0.14 **	1.60 ± 0.02 **
		25 μ M	0.52 ± 0.05 **	7.55 ± 0.56 **	1.77 ± 0.08 **	1.39 ± 0.06 **	0.87 ± 0.08 **	3.00 ± 0.09 **	0.79 ± 0.05 **	0.17 ± 0.005 **	1.34 ± 0.08 **	1.47 ± 0.08 **	0.97 ± 0.03 **

Results are given in μ M as mean ± SEM; *: $p < 0.01$, **: $p < 0.001$ by means of one-way ANOVA followed by Dunnett's post hoc test as compared to the IC₅₀ value of doxorubicin alone, $n = 3$. Values representing higher than two-fold sensitization to doxorubicin are highlighted in blue, those with higher than five-fold, in red.