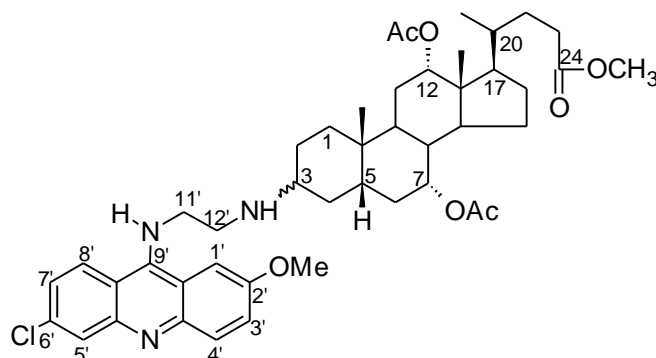


SUPPLEMENTARY MATERIAL TO
**New 9-aminoacridine derivatives as inhibitors of botulinum
neurotoxins and *P. falciparum* malaria**

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SPECTRAL AND ANALYTICAL DATA OF ALL THE SYNTHESIZED COMPOUNDS

Methyl 7 α ,12 α -diacetoxy-3 α -(N-{2-[(6-chloro-2-methoxyacridin-9-yl)amino]ethyl}amino)-5 β -cholan-24-oate (16). Yield: 9.6 %; Yellow foam, softens at 101–104 °C; ¹H-NMR (200 MHz, CDCl₃, δ / ppm): 8.12 (*d*, *J* = 9.0 Hz, H-C(8')), 8.05 (*d*, *J* = 2.2 Hz, H-C(5')), 7.98 (*d*, *J* = 9.6 Hz, H-C(4')), 7.41, (*dd*, *J*₁ = 2.4 Hz, *J*₂ = 9.0 Hz, H-C(3')), 7.45–7.25 (*m*, H-C(1'), H-C(7')), 5.10 (*bs*, H-C(12)), 4.95–4.85 (*m*, H-C(7')), 3.97 (*s*, ACR-OCH₃), 3.85–3.70 (*bs*, 2H-C(11')), 3.66 (*s*, CO₂CH₃), 3.05–2.90 (*m*, 2H-C(12')), 2.50–2.34 (*m*, H β -C(3)), 2.11 (*s*,

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CH₃COO), 1.98 (s, CH₃COO), 0.93 (s, CH₃-C(10)), 0.82 (d, $J = 6.2$ Hz, CH₃-C(20)), 0.74 (s, CH₃-C(13)); ¹³C-NMR (50 MHz CDCl₃, δ / ppm): 174.57, 170.57, 170.44, 155.91, 150.34, 148.17, 134.95, 131.13, 127.91, 124.45, 124.32, 117.93, 160.00, 99.58, 95.79, 75.56, 70.94, 57.76, 55.52, 51.51, 49.54, 47.38, 46.47, 45.10, 43.52, 41.66, 37.74, 37.31, 35.51, 34.80, 34.58, 31.48, 30.84, 30.75, 29.10, 28.95, 27.15, 25.62, 22.83, 21.52, 21.40, 17.46, 12.22; HRMS (m/z): 790.42058 corresponds to the molecular formula C₄₅H₆₀ClN₃O₇⁺ (error +1.67 ppm)

Methyl 7 α ,12 α -diacetoxo-3 β -(N-{2-[(6-chloro-2-methoxyacridin-9-yl)amino]ethyl}amino)-5 β -cholan-24-oate (17). Yield: 8.5 %; Yellow foam, softens at 102–105 °C; ¹H-NMR (500 MHz, CDCl₃, δ / ppm): 8.07 (d, $J = 9.3$ Hz, H-C(8')), 8.03 (d, $J = 2.0$ Hz, H-C(5')), 7.97 (d, $J = 9.3$ Hz, H-C(4')), 7.39 (dd, $J_1 = 2.6$ Hz, $J_2 = 9.4$ Hz, H-C(3')), 7.28 (d, $J = 2.6$ Hz, H-C(1')), 7.24 (dd, $J_1 = 2.1$ Hz, $J_2 = 9.2$ Hz, H-C(7')), 5.10 (bs, H-C(12)), 4.94–4.90 (m, H-C(7)), 3.94 (s, ACR-OCH₃), 3.79–3.74 (m, 2H-C(11')), 3.66 (s, CO₂CH₃), 2.93 (bs, H α -C(3)), 2.91–2.86 (m, 2H-C(12')), 2.39–2.30 (m, 1H), 2.25–2.15 (m, 3 \times H), 2.12 (s, CH₃COO), 2.08 (s, CH₃COO), 0.96 (s, CH₃-C(10)), 0.82 (d, $J = 6.6$ Hz, CH₃-C(20)), 0.74 (s, CH₃-C(13)); ¹³C-NMR (125 MHz, CDCl₃, δ / ppm): 174.50, 170.44, 170.34, 155.74, 150.37, 147.90, 135.0, 130.75, 127.56, 124.34, 124.09, 117.50, 115.54, 99.77, 75.47, 71.00, 55.45, 52.32, 51.49, 49.27, 47.34, 46.90, 45.03, 43.33, 37.72, 36.28, 34.91, 34.58, 33.39, 31.18, 30.87, 30.74, 30.17, 28.32, 27.15, 25.67, 25.02, 23.20, 22.79, 21.57, 21.31, 17.48, 12.17; HRMS (m/z): 790.41962 corresponds to the molecular formula C₄₅H₆₀ClN₃O₇H⁺ (error +0.46 ppm).

Methyl 7 α ,12 α -diacetoxo-3 β -(N-{3-[(6-chloro-2-methoxyacridin-9-yl)amino]propyl}amino)-5 β -cholan-24-oate (18). Yield: 15 %, contains ca. 20 % of the 3 α -isomer **19**; Yellow foam, softens at 100–103 °C; ¹H-NMR (500 MHz, CDCl₃, δ / ppm): 8.08 (d, $J = 9.6$ Hz, H-C(8')), 8.02 (d, $J = 1.8$ Hz, H-C(5')), 7.98 (d, $J = 9.6$, H-C(4')), 7.43–7.35 (m, N-H, H-C(3')), 7.36–7.25 (m, H-C(1')), 7.10 (dd, $J_1 = 2.2$ Hz, $J_2 = 9.20$ Hz, H-C(7')), 5.09 (s, H-C(12)), 4.91 (d, $J = 2.6$, H-C(7)), 3.95–3.90 (m, ACR-OCH₃, 2H-C(11')), 3.66 (s, CO₂CH₃), 2.93 (bs, H α -C(3)), 2.90–2.80 (m, 2H-C(13')), 2.38–2.30 (2H-C(24)), 2.11 (s, CH₃COO), 2.08 (s, CH₃COO), 0.87 (s, CH₃-C(10)), 0.82 (d, $J = 6.4$ Hz, CH₃-C(20)), 0.72 (s, CH₃-C(13)); ¹³C-NMR (125 MHz CDCl₃, δ / ppm): 174.53, 170.45, 170.38, 155.30, 150.73, 148.05, 140.51, 134.87, 130.95, 127.51, 124.51, 123.76, 123.66, 117.08, 115.10, 101.34, 75.51, 71.05, 55.59, 53.25, 51.51, 51.05, 47.36, 46.75, 45.05, 43.35, 37.72, 36.23, 34.88, 34.61, 33.16, 31.21, 30.89, 30.77, 30.17, 29.59, 28.46, 27.18, 25.68, 24.52, 23.04, 22.82, 21.60, 21.34, 17.51, 12.18; HRMS (m/z): 804.43541 corresponds to the molecular formula C₄₆H₆₂ClN₃O₇H⁺ (error +0.62 ppm).

Methyl 7 α ,12 α -diacetoxy-3 α -(N-{3-[(6-chloro-2-methoxyacridin-9-yl)amino]propyl}amino)-5 β -cholan-24-oate (19). Yield: 11 %, mixture of *ca.* 1:1.5 ratio of the β - and α -isomers; Yellow foam, softens at 100–103 °C; ¹H-NMR (500 MHz, CDCl₃, δ / ppm): 8.07–7.96 (*m*, H-C(8')), 7.73–7.65 (*m*, H-C(5'), H-C(4')), 7.42 (*bs*, H-C(1')), 7.24–7.11, (*m*, H-C(3')), H-C(7')), 5.10–5.03 (*m*, H-C(12)), 4.95–4.85 (*m*, H-C(7)), 4.17–4.09 (*bs*, 2H-C(11')), 3.91–3.86 (*m*, ACR-OCH₃), 3.65 (*s*, CO₂CH₃), 3.646 (*s*, CO₂CH₃), 3.33–3.19, (1H, *m*), 3.17–3.10 (*m*, 2H-C(12')), 3.07–2.99 (1H, *m*), 2.76–2.67 (1H, *m*), 2.10 (*s*, CH₃COO), 2.06 (*s*, CH₃COO), 0.94 (*s*, CH₃-C(10)), 0.83–0.77 (*m*, CH₃-C(20)), 0.73 (*s*, CH₃-C(13)), 0.72 (*s*, CH₃-C(13)); ¹³C-NMR (125 MHz CDCl₃, δ / ppm): 174.55, 170.99, 170.81, 155.95, 150.73, 148.05, 140.51, 134.87, 130.95, 127.51, 124.51, 123.76, 123.66, 117.08, 115.10, 101.34, 75.22, 70.54, 59.23, 56.05, 51.51, 47.38, 47.30, 45.10, 45.06, 43.33, 41.51, 37.85, 35.08, 34.71, 34.59, 31.61, 31.35, 30.87, 30.77, 29.65, 28.97, 27.17, 25.58, 22.83, 22.72, 21.55, 21.34, 21.14, 17.52, 17.46, 12.21.

3 α ,7 α ,12 α -Triacetoxy-24-{N-{2-[(6-chloro-2-methoxyacridin-9-yl)amino]ethyl}amino}-5 β -cholan (20). Yield: 42 %; Orange solid, m.p.: 115–117 °C; Anal. Calcd. for C₄₆H₆₂ClN₃O₇: C, 68.68; H, 7.77; N, 5.22 %. Found: C, 68.28; H, 7.52; N, 5.00 %; IR (ATR, cm⁻¹): 3324w, 2945m, 2870m, 1731s, 1632w, 1563m, 1522w, 1469m, 1375m, 1250s, 1068w, 1028m, 736w; ¹H-NMR (500 MHz, CDCl₃, δ / ppm): 8.08 (*d*, *J* = 9.0 Hz, H-C(8')), 8.01 (*d*, *J* = 2.0 Hz, H-C(5')), 7.96 (*d*, *J* = 9.0 Hz, H-C(4')), 7.37 (*dd*, *J*₁ = 2.0 Hz, *J*₂ = 9.0 Hz, H-C(3')), 7.32 (*d*, *J* = 2.5 Hz, H-C(1')), 7.23 (*dd*, *J*₁ = 2.5 Hz, *J*₂ = 9.0 Hz, H-C(3')), 5.11 (*bs*, H-C(12)), 4.92–4.90 (*m*, H-C(7)), 4.61–4.54 (*m*, H-C(3)), 3.95 (*s*, ACR-OCH₃), 3.84–3.80 (*m*, 2H-C(11')), 3.01–2.97 (*m*, 2H-C(12')), 2.72–2.63 (*m*, 2H-C(24)), 2.13 (*s*, CH₃COO-), 2.08 (*s*, CH₃COO-), 2.05 (*s*, CH₃COO-), 0.92 (*s*, H₃C-C(10)), 0.84 (*d*, *J* = 6.5 Hz, H₃C-C(20)), 0.74 (*s*, H₃C-C(13)); ¹³C-NMR (125 MHz, CDCl₃, δ / ppm): 170.50, 170.48, 170.34, 155.88, 150.78, 135.38, 124.75, 124.11, 117.44, 115.14, 99.77, 75.43, 74.07, 70.70, 55.50, 49.70, 49.03, 48.49, 47.67, 45.06, 43.40, 40.93, 37.75, 35.02, 34.69, 34.61, 34.33, 33.35, 31.24, 28.90, 27.33, 26.88, 26.74, 25.58, 22.82, 22.54, 21.59, 21.47, 21.43, 17.92, 14.17, 12.25; HRMS (*m/z*): 804.43437 corresponds to the molecular formula C₄₆H₆₂ClN₃O₇H⁺ (error -0.66 ppm), [M+2H]²⁺ = 402.72035 (error -1.85 ppm).

3 α ,7 α ,12 α -Triacetoxy-24-{N-{3-[(6-chloro-2-methoxyacridin-9-yl)amino]propyl}amino}-5 β -cholan (21). Yield: 29 %; Orange solid, m.p.: 93–94 °C; Anal. Calcd. for C₄₇H₆₄ClN₃O₇: C, 68.97; H, 7.88; N, 5.13 %. Found: C, 68.39; H, 7.60; N, 5.01 %; IR (ATR, cm⁻¹): 3258w, 2944m, 2869m, 1730s, 1632m, 1606w, 1565m, 1521m, 1469m, 1439m, 1375m, 1250s, 1119w, 1029m, 831w, 738w; ¹H-NMR (500 MHz, CDCl₃, δ / ppm): 8.10 (*d*, *J* = 9.3 Hz, H-C(8')), 8.01 (*d*, *J* = 2.1 Hz, H-C(5')), 7.96 (*d*, *J* = 9.3 Hz, H-C(4')), 7.39 (*dd*, *J*₁ =

= 2.7 Hz, $J_2 = 9.4$ Hz, H-C(3')), 7.35 (*d*, $J = 2.7$ Hz, H-C(1')), 7.22 (*dd*, $J_1 = 9.3$ Hz, $J_2 = 2.1$ Hz, H-C(7')), 5.09 (*bs*, H-C(12)), 4.92–4.88 (*m*, H-C(7)), 4.61–4.53 (*m*, H-C(3)), 3.98–3.91 (*m*, ACR-OCH₃ and 2H-C(11')), 2.94 (*t*, $J = 5.6$ Hz, 2H-C(13')), 2.73–2.61 (*m*, 2H-C(24)), 2.09 (*s*, CH₃COO⁻), 2.06 (*s*, CH₃COO⁻), 2.05 (*s*, CH₃COO⁻), 0.91 (*s*, H₃C-C(10)), 0.84 (*d*, $J = 6.6$ Hz, H₃C-C(20)), 0.72 (*s*, H₃C-C(13)), ¹³C-NMR (125 MHz, CDCl₃, δ / ppm): 170.52, 170.49, 170.36, 155.21, 150.61, 148.55, 146.80, 134.63, 131.14, 127.93, 124.73, 123.76, 123.45, 116.93, 114.93, 100.90, 75.42, 74.05, 70.67, 55.50, 51.10, 50.92, 49.35, 47.62, 45.04, 43.43, 40.91, 37.73, 35.00, 34.65, 34.59, 34.30, 33.41, 31.23, 30.11, 28.92, 27.27, 26.85, 26.53, 25.58, 22.78, 22.53, 21.56, 21.46, 21.38, 17.91, 12.24; HRMS (*m/z*): 818.44946 corresponds to the molecular formula C₄₇H₆₄ClN₃O₇H⁺ (error -1.34 ppm), 409.72839 corresponds to molecular formula C₄₇H₆₄ClN₃O₇H₂²⁺ (error -1.29 ppm).

N-[2-(1-Adamantyl)ethyl]-N'-(6-chloro-2-methoxyacridin-9-yl)ethane-1,2-diamine (**22**). Yield: 10 %; m.p.: 112–120 °C; IR (ATR, cm⁻¹): 3291*m*, 3061*w*, 2995*w*, 2908*s*, 2847*s*, 2654*w*, 1676*w*, 1632*m*, 1564*s*, 1527*s*, 1471*m*, 1343*w*, 1264*w*, 1240*m*, 1169*w*, 1123*w*, 1037*w*, 930*w*, 827*w*; ¹H-NMR (500 MHz, CDCl₃, δ / ppm): 8.09 (*d*, $J = 9.2$ Hz, H-C(8')), 8.03 (*d*, $J = 2.1$ Hz, H-C(5')), 7.97 (*d*, $J = 9.4$ Hz, H-C(4')), 7.39 (*dd*, $J_1 = 2.6$ Hz, $J_2 = 9.4$ Hz, H-C(3')), 7.32 (*d*, $J = 2.6$ Hz, H-C(1')), 7.25 (*dd*, $J_1 = 2.1$ Hz, $J_2 = 9.2$ Hz, H-C(7')), 3.96 (*s*, ACR-OCH₃), 3.78 (*t*, $J = 5.5$ Hz, 2H-C(11')), 2.96 (*t*, $J = 5.5$ Hz, 2H-C(12')), 2.73–2.67 (*m*, CH₂CH₂Ad), 1.95 (*bs*, 3H-Ad), 1.73–1.60 (*m*, 7H-Ad), 1.53–1.48 (*m*, 6H-Ad and NH), 1.40–1.33 (*m*, CH₂CH₂Ad); ¹³C-NMR (125 MHz, CDCl₃, δ / ppm): 155.89, 150.63, 124.76, 124.65, 124.13, 117.76, 99.52, 55.52, 49.33, 48.74, 45.01, 43.75, 42.71, 37.09, 31.91, 28.63; HRMS (*m/z*): 464.2458 corresponds to the molecular formula C₂₈H₃₄N₃ClOH⁺ (error: -1.08 ppm).

N-[2-(1-Adamantyl)ethyl]-N'-(6-chloro-2-methoxyacridin-9-yl)propane-1,3-diamine (**23**). Yield: 46 %; Brown–orange amorphous solid, m.p.: 76–78 °C; IR (ATR, cm⁻¹): 3270*m*, 3076*w*, 2903*s*, 2845*s*, 1732*w*, 1632*s*, 1566*s*, 1522*s*, 1446*s*, 1350*m*, 1241*s*, 1114*m*, 1073*w*, 1036*m*, 927*m*, 872*w*, 831*m*, 804*w*; ¹H-NMR (500 MHz, MeOD, δ / ppm): 8.25 (*d*, $J = 9.3$ Hz, H-C(8')), 7.82 (*d*, $J = 2.1$ Hz, H-C(5')), 7.79 (*d*, $J = 9.4$ Hz, H-C(4')), 7.52 (*d*, $J = 2.5$ Hz, H-C(1')), 7.42 (*dd*, $J_1 = 2.6$ Hz, $J_2 = 9.4$ Hz, H-C(3')), 7.28 (*dd*, $J_1 = 2.1$ Hz, $J_2 = 9.3$ Hz, H-C(7')), 3.97 (*s*, ACR-OCH₃), 3.92 (*t*, $J = 6.8$ Hz, 2H-C(11')), 2.80 (*t*, $J = 7.1$ Hz, 2H-C(13')), 2.63–2.59 (*m*, CH₂CH₂Ad), 2.02–1.96 (*m*, 2H-C(12')), 1.93 (*bs*, 3H-Ad and NH), 1.75–1.60 (*m*, 7H-Ad), 1.45–1.42 (*m*, 6H-Ad), 1.25–1.19 (*m*, CH₂CH₂Ad); ¹³C-NMR (125 MHz, MeOD, δ / ppm): 157.55; 153.23; 148.72; 137.01; 129.63; 127.48; 126.33; 126.17; 124.51; 118.79; 115.97; 101.97; 56.46; 47.90; 44.83; 43.64; 43.54; 38.22; 32.86; 30.54; 30.19; HRMS (*m/z*): 478.2603 corresponds to the molecular formula C₂₉H₃₆N₃ClOH⁺ (error: -3.58 ppm),

239.6341 corresponds to the molecular formula $C_{29}H_{36}N_3ClOH_2^{2+}$ (error -2.32 ppm).

N-Benzyl-N'-(6-chloro-2-methoxyacridin-9-yl)ethane-1,2-diamine (24). Yield: 60 %; Brown–orange amorphous solid, m.p.: 90–92°C; IR (ATR, cm^{-1}): 3366m, 3287m, 3066w, 3007w, 2963w, 2939w, 2899w, 2821m, 2709w, 1630m, 1606w, 1558s, 1524s, 1504s, 1437s, 1395w, 1339m, 1253s, 1234s, 1169m, 1141w, 1150m, 1082w, 1037m, 989w, 928m, 866w, 842w, 820m, 793w, 751s; 1H -NMR (500 MHz, $CDCl_3$): 8.06 (*d*, $J = 9.2$ Hz, H-C(8)), 8.04 (*d*, $J = 1.8$ Hz, H-C(5)), 7.97 (*d*, $J = 9.2$ Hz, H-C(4)), 7.39 (*dd*, $J_1 = 2.7$ Hz, $J_2 = 9.4$ Hz, H-C(3)), 7.36–7.33 (*m*, H-C(1) and 3H-Ar), 7.32–7.26 (*m*, 2H-Ar), 7.24 (*dd*, $J_1 = 2.1$ Hz, $J_2 = 9.2$ Hz, H-C(7)), 6.03 (*bs*, NH), 3.86 (*bs*, ArCH₂ and ACR-OCH₃), 3.75–3.72 (*m*, 2H-C(11)), 2.96–2.93 (*m*, 2H-C(12)), 1.71 (*bs*, NH); ^{13}C -NMR (125 MHz, $CDCl_3$, δ / ppm): 155.87, 150.15, 148.46, 146.75, 139.84, 134.65, 131.40, 128.56, 128.16, 128.03, 127.29, 124.48, 124.20, 118.06, 115.97, 99.23, 55.38, 53.36, 48.94, 48.80. HRMS (m/z): 392.1524 corresponds to the molecular formula $C_{23}H_{22}N_3ClOH^+$ (error: -1.57 ppm).

N-(6-Chloro-2-methoxyacridin-9-yl)ethane-1,2-diamine (12).¹ Yield: 89 %; 1H -NMR (500 MHz, CD_3OD , δ / ppm): 8.16 (*d*, $J = 9.2$ Hz, H-C(8)), 7.80 (*d*, $J = 2.1$ Hz, H-C(5)), 7.76 (*d*, $J = 9.4$ Hz, H-C(4)), 7.41 (*d*, $J = 2.6$ Hz, H-C(1)), 7.35 (*dd*, $J_1 = 2.2$ Hz, $J_2 = 9.3$ Hz, H-C(3)), 7.22 (*dd*, $J_1 = 2.2$ Hz, $J_2 = 9.3$ Hz, H-C(7)), 3.93 (*bs*, OCH₃), 3.77 (*t*, $J = 6.4$ Hz, 2H-C(11)), 2.99 (*t*, $J = 6.4$ Hz, 2H-C(12)); ^{13}C -NMR (125 MHz, CD_3OD , δ / ppm): 157.43, 152.80, 149.21, 147.00, 136.48, 130.46, 127.14, 126.98, 126.21, 124.55, 118.88, 116.25, 101.08, 56.28, 53.32, 43.38.

N-(6-Chloro-2-methoxyacridin-9-yl)propane-1,3-diamine (13).¹ Yield: 95 % 1H -NMR (200 MHz, $CDCl_3$, δ / ppm): 8.08–7.88 (*m*, H-C(8), H-C(5) and H-C(4)), 7.40–7.35 (*m*, H-C(3)), 7.28 (*d*, $J = 3.0$ Hz, H-C(1)), 7.14 (*dd*, $J_1 = 2.2$ Hz, $J_2 = 9.0$ Hz, H-C(7)), 3.93–3.83 (*m*, OCH₃, 2H-C(11)), 3.06 (*t*, $J = 5.6$ Hz, 2H-C(13)), 1.88–1.75 (*m*, 2H-C(12)); ^{13}C -NMR (50 MHz, $CDCl_3$, δ / ppm): 155.31, 150.39, 148.68, 146.17, 134.61, 130.77, 127.52, 125.27, 124.01, 123.01, 116.69, 114.24, 100.01, 55.33, 50.78, 41.37, 32.06.

N-(6-Chloro-2-methoxyacridin-9-yl)pentane-1,5-diamine (14).² Yield: 87 %; 1H -NMR (200 MHz, $CDCl_3$, δ / ppm): 8.04–7.90 (*m*, H-C(8), H-C(5) and H-C(4)), 7.43–7.35 (*m*, 1H), 7.29–7.10 (*m*, 2H), 4.72 (*bs*, NH), 3.93 (*bs*, OCH₃), 3.70–3.54 (*m*, 2H-C(11)), 2.68 (*bs*, 2H-C(15)), 1.85–1.60 (*m*, 2H-C(12)), 1.45 (*bs*, 2H-C(14) and 2H-C(13)), 1.22 (*bs*, NH₂); ^{13}C -NMR (50 MHz, $CDCl_3$, δ / ppm): 155.86, 149.57, 148.28, 146.70, 134.61, 131.40, 128.14, 124.36, 123.96, 117.78, 115.73, 98.99, 55.39, 50.49, 41.86, 33.18, 31.52, 24.09.

N-(6-Chloro-2-methoxyacridin-9-yl)hexane-1,6-diamine (15).³ Yield: 85 %; 1H -NMR (200 MHz, CD_3OD , δ / ppm): 8.07 (*d*, $J = 8.8$ Hz, H-C(8)), 7.79–7.71 (*m*, H-C(5) and H-C(4)), 7.36–7.28 (*m*, H-C(1) and H-C(3)), 7.15 (*dd*, $J_1 = 2.2$

Hz, $J_2 = 9.6$ Hz, H-C(7)), 3.90 (*s*, ACR-OCH₃), 3.69–3.61 (*m*, 2H-C(11)), 2.60–2.49 (*m*, 2H-C(16)), 1.78–1.61 (*m*, 2H-C(12)), 1.45–1.20 (*m*, 6H); ¹³C-NMR (50 MHz, CD₃OD, δ / ppm): 157.04, 152.69, 149.23, 146.90, 136.29, 130.35, 127.02, 126.95, 125.93, 124.02, 118.19, 115.59, 101.04, 56.10, 50.84, 42.37, 33.65, 32.12, 27.81, 27.68.

REFERENCES

1. S. Bongarzone, H. N. A. Tran, A. Cavalli, M. Roberti, P. Carloni, G. Legname and M. L. Bolognesi, *J. Med. Chem.* **53** (2010) 8197
2. C. Boulanger, C. Di Giorgio, P. Vierling, *Eur. J. Med. Chem.*, **40** (2005) 1295, described as triflate salt.
3. L. Guetzoyan, F. Ramiandrasoa, H. Dorizon, C. Desprez, A. Bridoux, C. Rogier, B. Pradines, M. Perrée-Fauveta, *Bioorg. Med. Chem.* **15** (2007) 3278.