Supplementary Figure 1:



Lack of a pro-inflammatory effect of PMA-Na on monocyte derived macrophages:

PMA (3.52 kDa) was incubated with monocyte-derived-macrophages for 3 h at 25 μ g/ml. It did not stimulate IFN- γ or TNF- α (n = 9). Similar results were obtained with 1.27 kDa, 1.7 kDa, 7.75 kDa and 18.5 kDa PMA.

Supplementary Figure 2:

UV spectra of AmB-PMA (3.52 kDa) and deoxycholate-AmB (Fungizone, Squibb).

The pale yellow powders were reconstituted in water at a final concentration of 10 µg/ml AmB.



AmB-PMA (3.52 kDa)

The AmB is highly aggregated in both deoxycholate-AmB (Fungizone, Squibb) and in liposomal AmB (Ambisome, Gilead). The λ_{max} of the UV spectra of deoxycholate-AmB was 329 nm (shown in red) and of liposomal AmB was 324 nm (not shown; see Reference 6). The λ_{max} of the UV spectra of AmB-PMA (3.52 kDa) (AP) was 324 nm (shown in blue). Similar results were obtained with 1.27 kDa, 1.7 kDa, 7.75 kDa and 18.5 kDa PMA.



Comparison of IRF-1 in BALB/c and C57BL/6 mouse lung at day +2:

Immuno-suppression caused a large and significant suppression of Interferon Regulatory Factor (IRF) -1 that did not change after infection, or with AF pre-exposure prophylaxis (n = 4).

For BALB/c mice: Normal IRF-1 = $1.24 \times 10^6 \pm 118,500$ copies/ 10^5 copies HPRT; IS + AF IRF-1 = $118,000 \pm 7,890$ copies/ 10^5 copies HPRT.

<u>For C57BL/6 mice:</u> Normal IRF-1 = $1.91 \times 10^6 \pm 210,800$ copies/ 10^5 copies HPRT; IS + AF IRF-1 = $157,300 \pm 16,200$ copies/ 10^5 copies HPRT.

Supplementary Table 1:

Isolate	Name	AmB- PMA	AmB- PMA	AmB- PMA	AmB- PMA	AmB- PMA	Fungizone
		(1.27 kDa)	(1.7 kDa)	(3.52 kDa)	(7.75 kDa)	(18.5 kDa)	
		MIC (µg/ml)	MIC (µg/ml)	MIC (µg/ml)	MIC (µg/ml)	MIC (µg/ml)	MIC (µg/ml)
J1	A. fumigatus	0.25	0.25	0.125	0.125	0.5	0.125
J2	A. fumigatus	0.25	0.25	0.125	0.125	0.25-0.5	0.25
J3	A. fumigatus	0.125	0.125	0.0625	0.0625	0.25	0.0625
J4	A. fumigatus	0.25	1.0	0.125	0.125	0.5	0.125-0.25
J5	A. fumigatus	0.25	1.0	0.125	0.125	0.5	0.125-0.25
HH-7	A. fumigatus	1.0	0.25	0.5	0.5	2.0	0.5
HH-12	A. fumigatus	0.5	0.25	0.25	0.25	1.0	0.25-0.5
HH-23	A. fumigatus	0.25	0.5	0.125	0.125	0.5	0.125-0.25
HH-24	A. fumigatus	0.5	0.5	0.25	0.25	1.0	0.25
HH-55	A. fumigatus	0.25	0.0625	0.125	0.125	0.25	0.125
CEA10	A. fumigatus	0.0625	0.25	0.0312	0.0312	0.125	0.0312

Summary of the MIC test results shown above:

Nanoparticle	Range of MICs (µg/ml)	Mean of MICs (µg/ml)
AmB-PMA (1.27 kDa)	0.0625 - 1	0.25
AmB-PMA (1.7 kDa)	0.0625 - 1	0.25
AmB-PMA (3.52 kDa)	0.0312 - 0.5	0.125
AmB-PMA (7.75 kDa)	0.0312 - 0.5	0.125
AmB-PMA (18.5 kDa)	0.125 - 2	0.5 - 1.0
Fungizone	0.0312 - 0.5	0.25

The antifungal activity of the different AmB-PMAs was determined against 11 isolates of *A*. *fumigatus* up to a final concentration of 16 μ g/ml. AmB-PMA (1.27 kDa & 1.7 kDa) had the same MIC as Fungizone (0.25 μ g/ml). In contrast, AmB-PMA (3.52 & 7.75 kDa) had better antifungal activity at 0.125 μ g/ml when compared to Fungizone. The anti-fungal activity of the AmB-PMA (18.5 kDa) was 0.5 - 1 μ g/ml.