Supplemental Online Content

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This supplemental material has been provided by the authors to give readers additional information about their work.

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eAppendix. Search Strategies

#1 ((clinical AND trial)[Title/Abstract] OR clinical trial[Title/Abstract] OR random*[Title/Abstract] OR random allocation[Title/Abstract])

#2 (“fungal infection*”[Title/Abstract] OR aspergillosis[Title/Abstract] OR blastomycosis[Title/Abstract] OR candidiasis[Title/Abstract] OR andidosis[Title/Abstract] OR coccidioidomycosis[Title/Abstract] OR cryptococcosis[Title/Abstract] OR histoplasmosis[Title/Abstract] OR mucormycosis[Title/Abstract] OR zygomycosis[Title/Abstract] OR paracoccidioidomycosis[Title/Abstract] OR “penicillium marneffei”[Title/Abstract] OR sporotrichosis[Title/Abstract] OR hyalohyphomycosis[Title/Abstract] OR fusariosis[Title/Abstract] OR fusarium[Title/Abstract] OR scedosporium[Title/Abstract] OR monosporium[Title/Abstract] OR phaeohyphomycosis[Title/Abstract] OR alternaria[Title/Abstract] OR bipolaris[Title/Abstract] OR curvularia[Title/Abstract] OR exserohilum[Title/Abstract] OR cladosiphialophora[Title/Abstract] OR xylorhpha[Title/Abstract] OR trichosporonosis[Title/Abstract] OR malassezia[Title/Abstract] OR pityrosporum[Title/Abstract] OR pneumocystis[Title/Abstract] OR pneumocystosis[Title/Abstract] OR pseudallescherias[Title/Abstract] OR allescherias[Title/Abstract] OR petriellidium[Title/Abstract])

#3 (antifungal[Title/Abstract] OR “antifungal agent*” OR amphotericin*[Title/Abstract] OR albaconazol*[Title/Abstract] OR UR9825[Title/Abstract] OR UR 9825[Title/Abstract] OR UR-9825[Title/Abstract] OR ketoconazol*[Title/Abstract] OR R41400[Title/Abstract] OR R 41400[Title/Abstract] OR R-41400[Title/Abstract] OR fluconazol*[Title/Abstract] OR UK49858[Title/Abstract] OR UK 49858[Title/Abstract] OR UK-49858[Title/Abstract] OR isavuconazol*[Title/Abstract] OR BAL8557[Title/Abstract] OR BAL 8557[Title/Abstract] OR BAL-8557[Title/Abstract] OR itraconazol*[Title/Abstract] OR R51211[Title/Abstract] OR R 51211[Title/Abstract] OR R-51211[Title/Abstract] OR posaconazol*[Title/Abstract] OR SCH56592[Title/Abstract] OR SCH 56592[Title/Abstract] OR SCH-56592[Title/Abstract] OR ravuconazol*[Title/Abstract] OR ER30346[Title/Abstract] OR ER 30346[Title/Abstract] OR ER-30346[Title/Abstract] OR voriconazol*[Title/Abstract] OR UK109496[Title/Abstract] OR UK 109496[Title/Abstract] OR UK-109496[Title/Abstract] OR HMR3270[Title/Abstract] OR HMR 3270[Title/Abstract] OR HMR-3270[Title/Abstract] OR IP960[Title/Abstract] OR IP 960[Title/Abstract] OR aminocandin*[Title/Abstract] OR anidulafungin*[Title/Abstract] OR LY303366[Title/Abstract] OR LY 303366[Title/Abstract] OR LY-303366[Title/Abstract] OR caspofungin*[Title/Abstract] OR MK0991[Title/Abstract] OR MK 0991[Title/Abstract] OR MK-0991[Title/Abstract] OR L743872[Title/Abstract] OR L 743872[Title/Abstract] OR L-743872[Title/Abstract] OR FK463[Title/Abstract] OR FK 463[Title/Abstract] OR flucytosin*[Title/Abstract] OR nystatin*[Title/Abstract])
**eFigure 1. PRISMA Flowchart**

- **Records identified through database searching**
  \( n = 35698 \)

- **Additional records identified through other sources**
  \( n = 11 \)

- **Records after duplicates removed**
  \( n = 1027 \)

- **Records screened**
  \( n = 1027 \)

- **Records excluded**
  \( n = 815 \)

- **Full-text articles assessed for eligibility**
  \( n = 212 \)

- **Studies included in qualitative synthesis**
  \( n = 69 \)

- **Studies included in quantitative synthesis (network meta-analysis)**
  \( n = 59 \)

*Reasons for exclusion:*
- Not appropriate study type \( (n=133) \)
- Not appropriate intervention \( (n=10) \)
eFigure 2. Quality Assessment
**eFigure 3.** Network Geometry of All Outcomes

Panel A. Fungal Infections

Legend: the blue nodes represent the interventions compared while the edges represent the direct comparisons available between pairs of interventions. The node sizes are weighted based on the number of patients in each arm while the edges are weighted using inverse variance. The edge colors (green or red) are weighted based on the quality of the studies included. A green edge indicates the comparison with high quality studies while a red line indicates comparison with low quality studies. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Legend: the blue nodes represent the interventions compared while the edges represent the direct comparisons available between pairs of interventions. The node sizes are weighted based on the number of patients in each arm while the edges are weighted using inverse variance. The edge colors (green or red) are weighted based on the quality of the studies included. A green edge indicates the comparison with high quality studies while a red line indicates comparison with low quality studies. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
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Panel D. Invasive Candidiasis

Legend: the blue nodes represent the interventions compared while the edges represent the direct comparisons available between pairs of interventions. The node sizes are weighted based on the number of patients in each arm while the edges are weighted using inverse variance. The edge colors (green or red) are weighted based on the quality of the studies included. A green edge indicates the comparison with high quality studies while a red line indicates comparison with low quality studies. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel E. Invasive Aspergillus

**Legend:** the blue nodes represent the interventions compared while the edges represent the direct comparisons available between pairs of interventions. The node sizes are weighted based on the number of patients in each arm while the edges are weighted using inverse variance. The edge colors (green or red) are weighted based on the quality of the studies included. A green edge indicates the comparison with high quality studies while a red line indicates comparison with low quality studies. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel F.  Mortality

Legend: the blue nodes represent the interventions compared while the edges represent the direct comparisons available between pairs of interventions. The node sizes are weighted based on the number of patients in each arm while the edges are weighted using inverse variance. The edge colors (green or red) are weighted based on the quality of the studies included. A green edge indicates the comparison with high quality studies while a red line indicates comparison with low quality studies. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel G. Fungi-related death

Legend: the blue nodes represent the interventions compared while the edges represent the direct comparisons available between pairs of interventions. The node sizes are weighted based on the number of patients in each arm while the edges are weighted using inverse variance. The edge colors (green or red) are weighted based on the quality of the studies included. A green edge indicates the comparison with high quality studies while a red line indicates comparison with low quality studies. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel H. Withdrawal

Legend: the blue nodes represent the interventions compared while the edges represent the direct comparisons available between pairs of interventions. The node sizes are weighted based on the number of patients in each arm while the edges are weighted using inverse variance. The edge colors (green or red) are weighted based on the quality of the studies included. A green edge indicates the comparison with high quality studies while a red line indicates comparison with low quality studies. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
eFigure 4. Forest Plots of All Outcomes

Panel A. Fungal Infections

Legend: The results are reported as Relative risks (RRs) and 95% confidence intervals (CIs). The blue line (line of null effect) is equal to 1. The solid black lines represent the CIs while the diamond summarizes the RRs. For each pairwise comparison, the forest plot should be read as following: if the diamond with the entire CI did not reach the blue line of null effect, there is a significant difference. If the entire CI is on the left of the null effect, the event is significantly higher in the “intervention arm” while, when the entire CI is on the right, the event is statistically more frequent in the “reference arm”. When the entire CI crosses the line of null effect, the difference between the two procedures compared is not statistically significant. In addition, a red line reports the Predictive Interval (Prl), namely the interval within which the estimate of a future study is expected to be. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel B. IFI

**Legend:** The results are reported as Relative risks (RRs) and 95% confidence intervals (CIs). The blue line (line of null effect) is equal to 1. The solid black lines represent the CIs while the diamond summarizes the RRs. For each pairwise comparison, the forest plot should be read as following: if the diamond with the entire CI did not reach the blue line of null effect, there is a significant difference. If the entire CI is on the left of the null effect, the event is significantly higher in the “intervention arm” while, when the entire CI is on the right, the event is statistically more frequent in the “reference arm”. When the entire CI crosses the line of null effect, the difference between the two procedures compared is not statistically significant. In addition, a red line reports the Predictive Interval (PrI), namely the interval within which the estimate of a future study is expected to be. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Legend: The results are reported as Relative risks (RRs) and 95% confidence intervals (CIs). The blue line (line of null effect) is equal to 1. The solid black lines represent the CIs while the diamond summarizes the RRs. For each pairwise comparison, the forest plot should be read as following: if the diamond with the entire CI did not reach the blue line of null effect, there is a significant difference. If the entire CI is on the left of the null effect, the event is significantly higher in the “intervention arm” while, when the entire CI is on the right, the event is statistically more frequent in the “reference arm”. When the entire CI crosses the line of null effect, the difference between the two procedures compared is not statistically significant. In addition, a red line reports the Predictive Interval (PrI), namely the interval within which the estimate of a future study is expected to be. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel D. Invasive Candidiasis

Legend: The results are reported as Relative risks (RRs) and 95% confidence intervals (CIs). The blue line (line of null effect) is equal to 1. The solid black lines represent the CIs while the diamond summarizes the RRs. For each pairwise comparison, the forest plot should be read as following: if the diamond with the entire CI did not reach the blue line of null effect, there is a significant difference. If the entire CI is on the left of the null effect, the event is significantly higher in the “intervention arm” while, when the entire CI is on the right, the event is statistically more frequent in the “reference arm”. When the entire CI crosses the line of null effect, the difference between the two procedures compared is not statistically significant. In addition, a red line reports the Predictive Interval (PrI), namely the interval within which the estimate of a future study is expected to be. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.

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Panel E. Invasive Aspergillus

Legend: The results are reported as Relative risks (RRs) and 95% confidence intervals (CIs). The blue line (line of null effect) is equal to 1. The solid black lines represent the CIs while the diamond summarizes the RRs. For each pairwise comparison, the forest plot should be read as following: if the diamond with the entire CI did not reach the blue line of null effect, there is a significant difference. If the entire CI is on the left of the null effect, the event is significantly higher in the “intervention arm” while, when the entire CI is on the right, the event is statistically more frequent in the “reference arm”. When the entire CI crosses the line of null effect, the difference between the two procedures compared is not statistically significant. In addition, a red line reports the Predictive Interval (Prl), namely the interval within which the estimate of a future study is expected to be. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
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Panel G. Fungi-related death

**Legend:** The results are reported as Relative risks (RRs) and 95% confidence intervals (CIs). The blue line (line of null effect) is equal to 1. The solid black lines represent the CIs while the diamond summarizes the RRs. For each pairwise comparison, the forest plot should be read as following: if the diamond with the entire CI did not reach the blue line of null effect, there is a significant difference. If the entire CI is on the left of the null effect, the event is significantly higher in the “intervention arm” while, when the entire CI is on the right, the event is statistically more frequent in the “reference arm”. When the entire CI crosses the line of null effect, the difference between the two procedures compared is not statistically significant. In addition, a red line reports the Predictive Interval (Prl), namely the interval within which the estimate of a future study is expected to be. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Legend: The results are reported as Relative risks (RRs) and 95% confidence intervals (CIs). The blue line (line of null effect) is equal to 1. The solid black lines represent the CIs while the diamond summarizes the RRs. For each pairwise comparison, the forest plot should be read as following: if the diamond with the entire CI did not reach the blue line of null effect, there is a significant difference. If the entire CI is on the left of the null effect, the event is significantly higher in the “intervention arm” while, when the entire CI is on the right, the event is statistically more frequent in the “reference arm”. When the entire CI crosses the line of null effect, the difference between the two procedures compared is not statistically significant. In addition, a red line reports the Predictive Interval (PrI), namely the interval within which the estimate of a future study is expected to be. AMB: conventional amphotericin B; AMLB: liposomal amphotericin B; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.

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Figure 5. League Tables of All Outcomes

Panel A. Fungal Infections

| Placebo       | MCFG     | CASP     | POCZ     | VOCZ     | ITZC     | KTCZ     | FLZC     | FLCZ     |
|---------------|----------|----------|----------|----------|----------|----------|----------|----------|
|               | 1.00     | 1.00     | 1.00     | 1.00     | 1.00     | 1.00     | 1.00     | 1.00     |
| 0.51          | 0.88     | 0.88     | 0.88     | 0.88     | 0.88     | 0.88     | 0.88     | 0.88     |
| (0.84, 0.95)  | (0.84, 0.95) | (0.84, 0.95) | (0.84, 0.95) | (0.84, 0.95) | (0.84, 0.95) | (0.84, 0.95) | (0.84, 0.95) | (0.84, 0.95) |
| 0.59          | 0.56     | 0.56     | 0.56     | 0.56     | 0.56     | 0.56     | 0.56     | 0.56     |
| (0.53, 0.64)  | (0.53, 0.64) | (0.53, 0.64) | (0.53, 0.64) | (0.53, 0.64) | (0.53, 0.64) | (0.53, 0.64) | (0.53, 0.64) | (0.53, 0.64) |
| 1.14          | 7.05     | 7.05     | 7.05     | 7.05     | 7.05     | 7.05     | 7.05     | 7.05     |
| (0.20, 2.83)  | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) |
| 1.04          | 4.99     | 4.99     | 4.99     | 4.99     | 4.99     | 4.99     | 4.99     | 4.99     |
| (0.20, 2.83)  | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) | (0.20, 2.83) |
| 0.42          | 2.81     | 2.81     | 2.81     | 2.81     | 2.81     | 2.81     | 2.81     | 2.81     |
| (0.30, 0.43)  | (0.30, 0.43) | (0.30, 0.43) | (0.30, 0.43) | (0.30, 0.43) | (0.30, 0.43) | (0.30, 0.43) | (0.30, 0.43) | (0.30, 0.43) |
| 3.51          | 3.51     | 3.51     | 3.51     | 3.51     | 3.51     | 3.51     | 3.51     | 3.51     |
| (0.33, 0.36)  | (0.33, 0.36) | (0.33, 0.36) | (0.33, 0.36) | (0.33, 0.36) | (0.33, 0.36) | (0.33, 0.36) | (0.33, 0.36) | (0.33, 0.36) |
| 0.23          | 0.94     | 0.94     | 0.94     | 0.94     | 0.94     | 0.94     | 0.94     | 0.94     |
| (0.28, 0.30)  | (0.28, 0.30) | (0.28, 0.30) | (0.28, 0.30) | (0.28, 0.30) | (0.28, 0.30) | (0.28, 0.30) | (0.28, 0.30) | (0.28, 0.30) |
| 0.65          | 2.07     | 2.07     | 2.07     | 2.07     | 2.07     | 2.07     | 2.07     | 2.07     |
| (0.30, 0.45)  | (0.30, 0.45) | (0.30, 0.45) | (0.30, 0.45) | (0.30, 0.45) | (0.30, 0.45) | (0.30, 0.45) | (0.30, 0.45) | (0.30, 0.45) |
| 0.69          | 2.35     | 2.35     | 2.35     | 2.35     | 2.35     | 2.35     | 2.35     | 2.35     |
| (0.43, 0.81)  | (0.43, 0.81) | (0.43, 0.81) | (0.43, 0.81) | (0.43, 0.81) | (0.43, 0.81) | (0.43, 0.81) | (0.43, 0.81) | (0.43, 0.81) |
| 0.71          | 4.18     | 4.18     | 4.18     | 4.18     | 4.18     | 4.18     | 4.18     | 4.18     |
| (0.25, 1.29)  | (0.25, 1.29) | (0.25, 1.29) | (0.25, 1.29) | (0.25, 1.29) | (0.25, 1.29) | (0.25, 1.29) | (0.25, 1.29) | (0.25, 1.29) |

Legend: RR >1 means the treatment in top left is better. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel B. IFI

| Placebo | MCFG | AMBL | AMB | Polyene |
|---------|------|------|-----|---------|
| 0.18    | 0.30 | 0.32 | 0.32 | 0.32    |
| (0.10, 0.50) | (0.12, 0.58) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |
| 0.87    | 0.32 | 0.30 | 0.30 | 0.30    |
| (0.52, 0.79) | (0.12, 0.14) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |
| 0.54    | 3.00 | 3.68 | 3.46 | 3.46    |
| (0.23, 1.70) | (0.22, 1.14) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |
| 0.02    | 2.01 | 3.02 | 0.19 | 0.98    |
| (0.14, 0.87) | (0.18, 1.10) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |
| 1.33    | 2.28 | 2.81 | 7.14 | 8.76    |
| (0.32, 0.73) | (0.46, 1.61) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |
| 1.08    | 1.39 | 6.62 | 14.01 | 14.01    |
| (0.33, 0.78) | (0.63, 0.72) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |
| 0.47    | 0.81 | 2.50 | 5.85 | 6.62    |
| (0.10, 0.28) | (0.20, 0.40) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |
| 0.44    | 1.74 | 2.16 | 5.50 | 5.60    |
| (0.26, 0.74) | (0.34, 0.22) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |
| 0.65    | 2.60 | 3.19 | 6.10 | 6.13    |
| (0.35, 0.70) | (0.48, 0.05) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |
| 1.10    | 3.14 | 6.57 | 14.11 | 14.11    |
| (0.18, 0.89) | (0.39, 0.29) | (0.01, 0.07) | (0.01, 0.24) | (0.01, 0.24) |

**Legend:** RR >1 means the treatment in top left is better. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel C. Proven IFI

Legend: RR >1 means the treatment in top left is better. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel D. Invasive Candidiasis

| Treatment | Placebo | MICG | CASP | POCZ | AMBL | KTCZ | ITCZ | FLNZ | MICZ | ITNZ |
|-----------|---------|------|------|------|------|------|------|------|------|------|
| RR        | 0.48    | 0.57 | 0.57 | 0.15 | 0.73 | 12.82| 0.59 | 0.18 | 0.33 | 1.73 |
| CI        | (0.02,1.11) | (0.10,1.42) | (0.19,1.83) | (0.09,1.62) | (0.04,1.33) | (2.55,3.03) | (2.55,3.03) | (0.81,1.00) | (0.06,1.00) | (0.57,3.30) |

**Legend:** RR >1 means the treatment in top left is better. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLNZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel E. Invasive Aspergillus

| Placebo (0.29±0.12) | MCFG (0.33±0.12) | CASP (0.36±0.17) | POCZ (0.29±0.12) | VOZ (0.36±0.17) | AMBL (0.39±0.12) | AMB (0.38±0.12) | Polyene (0.38±0.12) |
|----------------------|-------------------|------------------|-------------------|-----------------|-----------------|-----------------|-------------------|
| 1.22 (0.27±0.12)     |                   |                  |                   |                 |                 |                 |
| 0.71 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 0.23 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 1.47 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 2.33 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 0.21 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 0.99 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 2.17 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 1.39 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 1.36 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 0.38 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 0.95 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |
| 2.24 (0.95±0.12)     |                   |                  |                   |                 |                 |                 |

**Legend:** RR >1 means the treatment in top left is better. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel F. Mortality

Legend: RR >1 means the treatment in top left is better. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel G. Fungi-related death

|                | Placebo | MCFG | CASP | AMBL | AMB | Polyene |
|----------------|---------|------|------|------|-----|---------|
|                | 0.84 (0.80, 0.88) | 0.84 (0.80, 0.88) | 0.84 (0.80, 0.88) | 0.84 (0.80, 0.88) | 0.84 (0.80, 0.88) | 0.84 (0.80, 0.88) |
| 0.48 (0.41, 0.55) | 0.48 (0.41, 0.55) | 0.48 (0.41, 0.55) | 0.48 (0.41, 0.55) | 0.48 (0.41, 0.55) | 0.48 (0.41, 0.55) | 0.48 (0.41, 0.55) |
| 0.20 (0.18, 0.22) | 0.20 (0.18, 0.22) | 0.20 (0.18, 0.22) | 0.20 (0.18, 0.22) | 0.20 (0.18, 0.22) | 0.20 (0.18, 0.22) | 0.20 (0.18, 0.22) |
| 1.00 (0.88, 1.14) | 1.00 (0.88, 1.14) | 1.00 (0.88, 1.14) | 1.00 (0.88, 1.14) | 1.00 (0.88, 1.14) | 1.00 (0.88, 1.14) | 1.00 (0.88, 1.14) |
| 0.03 (0.01, 0.05) | 0.03 (0.01, 0.05) | 0.03 (0.01, 0.05) | 0.03 (0.01, 0.05) | 0.03 (0.01, 0.05) | 0.03 (0.01, 0.05) | 0.03 (0.01, 0.05) |
| 0.02 (0.01, 0.03) | 0.02 (0.01, 0.03) | 0.02 (0.01, 0.03) | 0.02 (0.01, 0.03) | 0.02 (0.01, 0.03) | 0.02 (0.01, 0.03) | 0.02 (0.01, 0.03) |
| 0.01 (0.00, 0.02) | 0.01 (0.00, 0.02) | 0.01 (0.00, 0.02) | 0.01 (0.00, 0.02) | 0.01 (0.00, 0.02) | 0.01 (0.00, 0.02) | 0.01 (0.00, 0.02) |
| 0.00 (0.00, 0.00) | 0.00 (0.00, 0.00) | 0.00 (0.00, 0.00) | 0.00 (0.00, 0.00) | 0.00 (0.00, 0.00) | 0.00 (0.00, 0.00) | 0.00 (0.00, 0.00) |

**Legend:** RR >1 means the treatment in top left is better. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
Panel H. Withdrawal

| Placebo | MCFG | CASP | AMB | AMBL | MICZ | KTCZ | Polyene |
|---------|------|------|-----|------|------|------|---------|
| 1.20    | 0.81 | 0.88 | 0.68| 0.68 | 0.62 | 0.39 | 0.86    |
| (0.34,2.50) | (0.55,7.52) | (0.19,4.91) | (0.01,6.04) | (0.62,8.84) | (0.02,1.02) | (0.02,1.23) | (0.20,3.92) |
| 0.22    | 2.83 | 2.35 | 0.28| 3.43 | 0.56 | 0.49 | 0.04    |
| (0.26,2.59) | (0.71,11.25) | (0.05,1.58) | (0.80,14.73) | (0.10,1.81) | (0.10,2.51) | (0.00,0.83) | (0.00,0.83) |
| 0.68    | 0.88 | 0.68 | 0.68| 0.62 | 0.39 | 0.39 | 0.90    |
| (0.26,2.59) | (0.62,8.74) | (0.01,6.04) | (0.02,1.02) | (0.02,1.23) | (0.02,1.23) | (0.02,1.23) | (0.20,3.92) |
| 0.88    | 0.88 | 0.62 | 0.56| 0.49 | 0.49 | 0.04 | 0.04    |
| (0.62,8.74) | (0.80,14.73) | (0.10,1.81) | (0.10,2.51) | (0.10,2.51) | (0.00,0.83) | (0.00,0.83) | (0.00,0.83) |
| 2.83    | 1.59 | 1.32 | 0.56| 1.89 | 0.99 | 0.04 | 0.04    |
| (0.71,11.25) | (0.76,3.31) | (0.05,1.58) | (0.10,1.81) | (0.10,2.51) | (0.00,0.83) | (0.00,0.83) | (0.00,0.83) |
| 2.35    | 1.59 | 1.32 | 0.56| 1.89 | 0.99 | 0.04 | 0.04    |
| (0.62,8.74) | (0.76,3.31) | (0.05,1.58) | (0.10,1.81) | (0.10,2.51) | (0.00,0.83) | (0.00,0.83) | (0.00,0.83) |
| 0.28    | 0.56 | 0.49 | 0.49| 0.04 | 0.04 | 0.04 | 0.04    |
| (0.05,1.58) | (0.10,1.81) | (0.00,0.83) | (0.00,0.83) | (0.00,0.83) | (0.00,0.83) | (0.00,0.83) | (0.00,0.83) |
| 2.83    | 2.35 | 2.35 | 2.35| 2.35| 2.35| 2.35| 2.35    |
| (0.62,8.74) | (0.80,14.73) | (0.80,14.73) | (0.80,14.73) | (0.80,14.73) | (0.80,14.73) | (0.80,14.73) | (0.80,14.73) |

Legend: RR >1 means the treatment in top left is better. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; MICZ: miconazole; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
**eFigure 6. Cluster Rank Combined the SUCRA**

Legend: The y axis reports the SUCRA values as a percentage of “Withdrawal”. The x axis reports the IFI. Different colors identify the different clusters. AMB: conventional amphotericin B; AMBL: liposomal amphotericin B; KTCZ: ketoconazole; FLCZ: fluconazole; ITCZ: itraconazole; VOCZ: voriconazole; POCZ: posaconazole; CASP: caspofungin; MCFG: micafungin.
**eFigure 7. Inconsistency Plot of All Outcomes**

Panel A. Fungal Infections

| Loop       | IF       | 95%CI (truncated) | Loop-specific Heterogeneity($I^2$) |
|------------|----------|-------------------|-----------------------------------|
| 08-10-12   | 1.40     | (0.00, 2.92)      | 0.000                             |
| 01-08-12   | 1.33     | (0.00, 3.28)      | 0.036                             |
| 01-03-06   | 1.15     | (0.00, 2.31)      | 0.000                             |
| 07-08-11   | 0.98     | (0.00, 2.44)      | 0.000                             |
| 07-08-10   | 0.97     | (0.00, 2.12)      | 0.000                             |
| 04-07-10   | 0.95     | (0.00, 4.34)      | 0.000                             |
| 01-07-09   | 0.92     | (0.00, 2.92)      | 0.179                             |
| 01-04-10-12| 0.67     | (0.00, 5.12)      | 0.659                             |
| 01-07-08   | 0.63     | (0.00, 1.27)      | 0.112                             |
| 01-08-09   | 0.63     | (0.00, 2.54)      | 0.000                             |
| 01-04-08-10| 0.57     | (0.00, 4.25)      | 0.344                             |
| 07-08-12   | 0.41     | (0.00, 1.37)      | 0.000                             |
| 01-03-08   | 0.41     | (0.00, 0.97)      | 0.006                             |
| 07-08-09   | 0.37     | (0.00, 1.66)      | 0.000                             |
| 01-07-12   | 0.29     | (0.00, 2.05)      | 0.144                             |
| 03-07-08   | 0.15     | (0.00, 0.57)      | 0.000                             |
| 01-04-07   | 0.10     | (0.00, 2.10)      | 0.247                             |
| 01-03-07   | 0.06     | (0.00, 0.85)      | 0.134                             |
| 07-10-12   | 0.05     | (0.00, 0.82)      | 0.000                             |

**Legend:** Inconsistency plot assuming loop-specific heterogeneity estimates using the method of moments estimator. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel B. IFI

Legend: Inconsistency plot assuming loop-specific heterogeneity estimates using the method of moments estimator. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel C. Proven IFI

Legend: Inconsistency plot assuming loop-specific heterogeneity estimates using the method of moments estimator. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel D. Invasive Candidiasis

| Loop   | 95%CI       | Loop-specific |
|--------|-------------|---------------|
|        | IF (truncated) | Heterogeneity($\chi^2$) |
| 01-03-07 | (1.07, 3.98) | 0.000         |
| 01-03-08 | (0.00, 2.90) | 0.000         |
| 07-08-12 | (0.00, 4.38) | 0.000         |
| 01-07-08 | (0.00, 2.31) | 0.000         |
| 03-07-08 | (0.00, 1.87) | 0.000         |
| 07-08-11 | (0.00, 2.76) | 0.000         |
| 01-04-07 | (0.00, 3.19) | 0.000         |
| 07-08-10 | (0.00, 3.32) | 0.000         |

**Legend:** Inconsistency plot assuming loop-specific heterogeneity estimates using the method of moments estimator. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel E. Invasive Aspergillus

Legend: Inconsistency plot assuming loop-specific heterogeneity estimates using the method of moments estimator. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel F. Mortality

Legend: Inconsistency plot assuming loop-specific heterogeneity estimates using the method of moments estimator. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel G. Fungi-related death

Legend: Inconsistency plot assuming loop-specific heterogeneity estimates using the method of moments estimator. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel H. Withdrawal

Legend: Inconsistency plot assuming loop-specific heterogeneity estimates using the method of moments estimator. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASp); arm 12= micafungin (MCFG).
eFigure 8. Funnel Plots of All Outcomes

Panel A. Fungal Infections

Legend: In the comparison-adjusted funnel plot, the horizontal axis shows the difference of each i-study estimate YiXY from the summary effect for the respective comparison (YiXY-μXY) while the vertical axis presents the measure of dispersion of YiXY, namely the standard error of the effect size. The red line shows the null hypothesis. Each point represents a direct comparison; different colors correspond to different comparisons. The dashed black line represents the 95% confidence interval. The horizontal line represents the regression line; the regression line demonstrates that no asymmetry is present. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Legend: In the comparison-adjusted funnel plot, the horizontal axis shows the difference of each i-study estimate YiXY from the summary effect for the respective comparison (YiXY-µXY) while the vertical axis presents the measure of dispersion of YiXY, namely the standard error of the effect size. The red line shows the null hypothesis. Each point represents a direct comparison; different colors correspond to different comparisons. The dashed black line represents the 95% confidence interval. The horizontal line represents the regression line; the regression line demonstrates that no asymmetry is present. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Legend: In the comparison-adjusted funnel plot, the horizontal axis shows the difference of each i-study estimate YiXY from the summary effect for the respective comparison (YiXY−μXY) while the vertical axis presents the measure of dispersion of YiXY, namely the standard error of the effect size. The red line shows the null hypothesis. Each point represents a direct comparison; different colors correspond to different comparisons. The dashed black line represents the 95% confidence interval. The horizontal line represents the regression line; the regression line demonstrates that no asymmetry is present. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel D. Invasive Candidiasis

Legend: In the comparison-adjusted funnel plot, the horizontal axis shows the difference of each i-study estimate $Y_{iXY}$ from the summary effect for the respective comparison ($Y_{iXY}-\mu_{XY}$) while the vertical axis presents the measure of dispersion of $Y_{iXY}$, namely the standard error of the effect size. The red line shows the null hypothesis. Each point represents a direct comparison; different colors correspond to different comparisons. The dashed black line represents the 95% confidence interval. The horizontal line represents the regression line; the regression line demonstrates that no asymmetry is present. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel E. Invasive Aspergillus

Legend: In the comparison-adjusted funnel plot, the horizontal axis shows the difference of each i-study estimate $Y_{iXY}$ from the summary effect for the respective comparison ($Y_{iXY} - \mu_{XY}$) while the vertical axis presents the measure of dispersion of $Y_{iXY}$, namely the standard error of the effect size. The red line shows the null hypothesis. Each point represents a direct comparison; different colors correspond to different comparisons. The dashed black line represents the 95% confidence interval. The horizontal line represents the regression line; the regression line demonstrates that no asymmetry is present. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel F. Mortality

Legend: In the comparison-adjusted funnel plot, the horizontal axis shows the difference of each i-study estimate YiXY from the summary effect for the respective comparison (YiXY-μXY) while the vertical axis presents the measure of dispersion of YiXY, namely the standard error of the effect size. The red line shows the null hypothesis. Each point represents a direct comparison; different colors correspond to different comparisons. The dashed black line represents the 95% confidence interval. The horizontal line represents the regression line; the regression line demonstrates that no asymmetry is present. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel G. Fungi-related death

Legend: In the comparison-adjusted funnel plot, the horizontal axis shows the difference of each i-study estimate $Y_{iXY}$ from the summary effect for the respective comparison ($Y_{iXY} - \mu_{XY}$) while the vertical axis presents the measure of dispersion of $Y_{iXY}$, namely the standard error of the effect size. The red line shows the null hypothesis. Each point represents a direct comparison; different colors correspond to different comparisons. The dashed black line represents the 95% confidence interval. The horizontal line represents the regression line; the regression line demonstrates that no asymmetry is present. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).
Panel H. Withdrawal

Legend: In the comparison-adjusted funnel plot, the horizontal axis shows the difference of each i-study estimate $Y_i^{XY}$ from the summary effect for the respective comparison ($Y_i^{XY} - \mu_{XY}$) while the vertical axis presents the measure of dispersion of $Y_i^{XY}$, namely the standard error of the effect size. The red line shows the null hypothesis. Each point represents a direct comparison; different colors correspond to different comparisons. The dashed black line represents the 95% confidence interval. The horizontal line represents the regression line; the regression line demonstrates that no asymmetry is present. arm 01= placebo; arm 02= oral polyene; arm 03= conventional amphotericin B (AMB); arm 04= liposomal amphotericin B (AMBL); arm 05= miconazole (MICZ); arm 06= ketoconazole (KTCZ); arm 07= fluconazole (FLCZ); arm 08= itraconazole (ITCZ); arm 09= voriconazole (VOCZ); arm 10= posaconazole (POCZ); arm 11= caspofungin (CASP); arm 12= micafungin (MCFG).