Supplement 6: Standard meta-analysis forest plot for a) active tuberculosis and b) hepatotoxicity

a)

**No treatment vs. INH 12-72 months**

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
<th>Weight %</th>
</tr>
</thead>
<tbody>
<tr>
<td>Agarwal</td>
<td>2004</td>
<td>0.36 (0.09, 1.36)</td>
<td>19.15</td>
<td>1.00878</td>
</tr>
<tr>
<td>Fitzgerald</td>
<td>2001</td>
<td>1.34 (0.37, 4.87)</td>
<td>19.97</td>
<td>11.4</td>
</tr>
<tr>
<td>Naqvi</td>
<td>2010</td>
<td>0.07 (0.01, 0.51)</td>
<td>10.56</td>
<td>1</td>
</tr>
<tr>
<td>Pape</td>
<td>1993</td>
<td>0.33 (0.10, 1.10)</td>
<td>21.61</td>
<td>114</td>
</tr>
<tr>
<td>Vikrant</td>
<td>2005</td>
<td>0.41 (0.17, 1.02)</td>
<td>28.72</td>
<td>1</td>
</tr>
<tr>
<td>D+L Overall (I-squared = 40.3%, p = 0.153)</td>
<td></td>
<td>0.40 (0.19, 0.84)</td>
<td>100.00</td>
<td>1</td>
</tr>
<tr>
<td>M-H Overall</td>
<td></td>
<td>0.36 (0.22, 0.61)</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

NOTE: Weights are from random effects analysis
### Placebo vs. INH 3-4 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>IUATCP</td>
<td>1982</td>
<td>0.78 (0.58, 1.06)</td>
<td>59.30</td>
</tr>
<tr>
<td>Veening</td>
<td>1968</td>
<td>0.07 (0.01, 0.67)</td>
<td>40.70</td>
</tr>
<tr>
<td>D+L Overall</td>
<td></td>
<td>0.30 (0.03, 3.03)</td>
<td>100.00</td>
</tr>
<tr>
<td>M-H Overall</td>
<td></td>
<td>0.70 (0.53, 0.95)</td>
<td></td>
</tr>
</tbody>
</table>

NOTE: Weights are from random effects analysis

### Placebo vs. INH 6 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gordin</td>
<td>1997</td>
<td>0.49 (0.12, 1.97)</td>
<td>2.56</td>
</tr>
<tr>
<td>Hawken</td>
<td>1997</td>
<td>1.09 (0.61, 1.97)</td>
<td>16.22</td>
</tr>
<tr>
<td>Horwitz</td>
<td>1966</td>
<td>0.67 (0.36, 1.28)</td>
<td>24.77</td>
</tr>
<tr>
<td>Johnson/Whalen (PPD positive)</td>
<td>2001</td>
<td>0.68 (0.43, 1.09)</td>
<td>13.29</td>
</tr>
<tr>
<td>Johnson/Whalen (PPD anergic)</td>
<td>2001</td>
<td>0.62 (0.32, 1.18)</td>
<td>8.91</td>
</tr>
<tr>
<td>Metenga/Quigley</td>
<td>1998</td>
<td>0.70 (0.42, 1.15)</td>
<td>12.34</td>
</tr>
<tr>
<td>Xie</td>
<td>2009</td>
<td>0.15 (0.02, 1.19)</td>
<td>1.21</td>
</tr>
<tr>
<td>IUATCP</td>
<td>1982</td>
<td>0.35 (0.24, 0.52)</td>
<td>15.89</td>
</tr>
<tr>
<td>HKC/STBCBMRC</td>
<td>1992</td>
<td>0.61 (0.35, 1.08)</td>
<td>10.81</td>
</tr>
<tr>
<td>D+L Overall</td>
<td></td>
<td>0.67 (0.48, 0.97)</td>
<td>100.00</td>
</tr>
<tr>
<td>M-H Overall</td>
<td></td>
<td>0.62 (0.55, 0.71)</td>
<td></td>
</tr>
</tbody>
</table>

NOTE: Weights are from random effects analysis
### Placebo vs. INH 12-72 months

**Author** | **Year** | **OR (95% CI)** | **Weight**
--- | --- | --- | ---
John | 1994 | 0.68 (0.29, 1.62) | 5.50
Madhi (HIV) | 2011 | 0.80 (0.48, 1.32) | 9.02
Madhi (not HIV) | 2011 | 0.89 (0.52, 1.52) | 8.75
Mohammed | 2007 | 1.69 (0.55, 5.18) | 3.96
Zar | 2007 | 0.36 (0.12, 1.03) | 4.26
IUATCP | 1982 | 0.25 (0.16, 0.39) | 9.73
Ferebee | 1962 | 0.43 (0.32, 0.59) | 11.36
Mount | 1962 | 0.46 (0.17, 1.23) | 4.72
Ferebee | 1963 | 0.36 (0.21, 0.60) | 8.79
Del Castillo | 1965 | 0.96 (0.48, 1.91) | 7.04
Bush | 1965 | 0.70 (0.28, 1.74) | 5.17
Comstock | 1967 | 0.40 (0.29, 0.54) | 11.38
Egsmose | 1965 | 0.17 (0.04, 0.76) | 2.50
Falk | 1978 | 0.70 (0.38, 1.29) | 7.62

**D+L Overall** (I-squared = 62.0%, p = 0.001)

**M-H Overall**

**NOTE:** Weights are from random effects analysis

### Placebo vs. RMP-INH 3-4 months

**Author** | **Year** | **OR (95% CI)** | **Weight**
--- | --- | --- | ---
Johnson/Whalen (PPD positive) | 2001 | 0.41 (0.24, 0.70) | 51.60
HKCSTBCBMRC | 1992 | 0.67 (0.38, 1.17) | 48.40

**D+L Overall** (I-squared = 33.8%, p = 0.219)

**M-H Overall**

**NOTE:** Weights are from random effects analysis
### Placebo vs. RMP-INH-PZA

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Johnson/Whalen (PPD positive)</td>
<td>2001</td>
<td>0.34 (0.18, 0.62)</td>
<td>56.46</td>
</tr>
<tr>
<td>Cowie</td>
<td>1996</td>
<td>0.72 (0.32, 1.60)</td>
<td>43.54</td>
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<tr>
<td>D+L Overall (I-squared = 53.7%, p = 0.141)</td>
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<td>0.47 (0.22, 0.98)</td>
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<tr>
<td>M-H Overall</td>
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<td>0.44 (0.27, 0.70)</td>
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</table>

**NOTE:** Weights are from random effects analysis

### INH 6 vs. INH 12-72 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Martinson</td>
<td>2011</td>
<td>0.71 (0.31, 1.63)</td>
<td>17.44</td>
</tr>
<tr>
<td>Samandari</td>
<td>2011</td>
<td>0.57 (0.33, 1.00)</td>
<td>38.55</td>
</tr>
<tr>
<td>IUATCP</td>
<td>1982</td>
<td>0.71 (0.42, 1.20)</td>
<td>44.01</td>
</tr>
<tr>
<td>D+L Overall (I-squared = 0.0%, p = 0.833)</td>
<td></td>
<td>0.65 (0.46, 0.92)</td>
<td>100.00</td>
</tr>
<tr>
<td>M-H Overall</td>
<td></td>
<td>0.65 (0.46, 0.92)</td>
<td></td>
</tr>
</tbody>
</table>

**NOTE:** Weights are from random effects analysis
INH 6 vs. RMP-INH 3-4 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Geijo</td>
<td>2007</td>
<td>0.29 (0.01, 7.25)</td>
<td>0.99</td>
</tr>
<tr>
<td>Johnson/Whalen (PPD positive)</td>
<td>2001</td>
<td>0.61 (0.35, 1.05)</td>
<td>34.14</td>
</tr>
<tr>
<td>Martinson</td>
<td>2011</td>
<td>1.09 (0.60, 1.99)</td>
<td>28.69</td>
</tr>
<tr>
<td>Rivero</td>
<td>2007</td>
<td>1.36 (0.35, 5.20)</td>
<td>5.72</td>
</tr>
<tr>
<td>JHC/STBC/MBRC</td>
<td>1992</td>
<td>1.09 (0.60, 1.98)</td>
<td>23.12</td>
</tr>
<tr>
<td>Jimenez-Fuentes</td>
<td>2013</td>
<td>0.39 (0.06, 15.95)</td>
<td>1.34</td>
</tr>
</tbody>
</table>

D+L Overall (I-squared = 0.0%, p = 0.610)

M-H Overall

NOTE: Weights are from random effects analysis

INH 6 vs. RMP-PZA

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
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<tbody>
<tr>
<td>Halsey</td>
<td>1998</td>
<td>0.97 (0.38, 2.48)</td>
<td>21.81</td>
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<tr>
<td>Magdor</td>
<td>1994</td>
<td>3.06 (0.12, 76.95)</td>
<td>1.83</td>
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<tr>
<td>Mwinga/Quigley</td>
<td>1998</td>
<td>1.16 (0.69, 1.95)</td>
<td>69.91</td>
</tr>
<tr>
<td>Rivero</td>
<td>2007</td>
<td>0.52 (0.09, 2.90)</td>
<td>6.45</td>
</tr>
</tbody>
</table>

D+L Overall (I-squared = 0.0%, p = 0.751)

M-H Overall

NOTE: Weights are from random effects analysis
INH 12-72 vs. RMP-INH 3-4 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Martinez Alfaro</td>
<td>2000</td>
<td>0.45 (0.08, 2.53)</td>
<td>30.05</td>
</tr>
<tr>
<td>Martinson</td>
<td>2011</td>
<td>1.53 (0.67, 3.49)</td>
<td>69.95</td>
</tr>
<tr>
<td>D+L Overall</td>
<td></td>
<td>1.06 (0.35, 3.21)</td>
<td></td>
</tr>
<tr>
<td>M-H Overall</td>
<td></td>
<td>1.22 (0.60, 2.50)</td>
<td></td>
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</table>

NOTE: Weights are from random effects analysis

RMP-INH 3-4 months vs. RMP-INH-PZA

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gupta</td>
<td>1993</td>
<td>0.11 (0.01, 2.12)</td>
<td>23.62</td>
</tr>
<tr>
<td>Johnson/Whalen (PPD positive)</td>
<td>2001</td>
<td>0.81 (0.42, 1.59)</td>
<td>76.38</td>
</tr>
<tr>
<td>D+L Overall</td>
<td></td>
<td>0.51 (0.10, 2.72)</td>
<td>100.00</td>
</tr>
<tr>
<td>M-H Overall</td>
<td></td>
<td>0.69 (0.36, 1.30)</td>
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</tbody>
</table>

NOTE: Weights are from random effects analysis
### No treatment vs. INH 12-72 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Bailey</td>
<td>1974</td>
<td>26.01 (1.50, 451.04)</td>
<td>40.14</td>
</tr>
<tr>
<td>Vikrant</td>
<td>2005</td>
<td>1.63 (0.54, 4.95)</td>
<td>59.86</td>
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</tbody>
</table>

**D+L Overall** (I-squared = 73.3%, p = 0.053)

**M-H Overall**

**Note:** Weights are from random effects analysis

### Placebo vs. INH 12-72 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>John</td>
<td>1994</td>
<td>1.05 (0.57, 1.92)</td>
<td>54.55</td>
</tr>
<tr>
<td>Madhi (HIV)</td>
<td>2011</td>
<td>0.20 (0.02, 1.70)</td>
<td>15.59</td>
</tr>
<tr>
<td>Madhi (not HIV)</td>
<td>2011</td>
<td>0.37 (0.10, 1.40)</td>
<td>29.86</td>
</tr>
</tbody>
</table>

**D+L Overall** (I-squared = 46.3%, p = 0.156)

**M-H Overall**

**Note:** Weights are from random effects analysis
### INH 6 vs. INH 12-72 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
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<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Martinson</td>
<td>2011</td>
<td>4.95 (2.68, 9.15)</td>
<td>56.21</td>
</tr>
<tr>
<td>Samandari</td>
<td>2011</td>
<td>1.48 (0.52, 4.17)</td>
<td>43.79</td>
</tr>
<tr>
<td>D+L Overall</td>
<td></td>
<td>2.92 (0.90, 9.44)</td>
<td>100.00</td>
</tr>
<tr>
<td>M-H Overall</td>
<td></td>
<td>3.55 (2.10, 6.01)</td>
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</tr>
</tbody>
</table>

NOTE: Weights are from random effects analysis

### INH 6 vs. RMP-INH 3-4 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Geijo</td>
<td>2007</td>
<td>0.88 (0.17, 4.57)</td>
<td>11.14</td>
</tr>
<tr>
<td>Martinson</td>
<td>2011</td>
<td>0.87 (0.43, 1.78)</td>
<td>60.05</td>
</tr>
<tr>
<td>Rivero</td>
<td>2007</td>
<td>1.08 (0.36, 3.18)</td>
<td>25.85</td>
</tr>
<tr>
<td>Jimenez-Fuentes</td>
<td>2013</td>
<td>0.33 (0.01, 8.13)</td>
<td>2.96</td>
</tr>
<tr>
<td>D+L Overall</td>
<td></td>
<td>0.89 (0.52, 1.55)</td>
<td>100.00</td>
</tr>
<tr>
<td>M-H Overall</td>
<td></td>
<td>0.89 (0.51, 1.54)</td>
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</tr>
</tbody>
</table>

NOTE: Weights are from random effects analysis
INH 6 vs. RMP-PZA

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L) Weight</th>
</tr>
</thead>
<tbody>
<tr>
<td>Leung</td>
<td>2003</td>
<td>39.94 (2.28, 699.73)</td>
<td>8.10</td>
</tr>
<tr>
<td>Rivero</td>
<td>2007</td>
<td>1.93 (0.73, 5.11)</td>
<td>37.26</td>
</tr>
<tr>
<td>Tortajada</td>
<td>2005</td>
<td>4.22 (1.50, 11.88)</td>
<td>35.14</td>
</tr>
<tr>
<td>Sanchez-Arcilla</td>
<td>2004</td>
<td>2.72 (0.51, 14.43)</td>
<td>19.51</td>
</tr>
</tbody>
</table>

D+L Overall: (I-squared = 36.3%, p = 0.195)

M-H Overall: 3.47 (1.46, 8.25)

NOTE: Weights are from random effects analysis

INH 9 months vs. RMP

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L) Weight</th>
</tr>
</thead>
<tbody>
<tr>
<td>Menzies</td>
<td>2004</td>
<td>0.14 (0.01, 2.68)</td>
<td>11.41</td>
</tr>
<tr>
<td>Menzies</td>
<td>2008</td>
<td>0.18 (0.05, 0.63)</td>
<td>66.10</td>
</tr>
<tr>
<td>White</td>
<td>2012</td>
<td>0.17 (0.02, 1.39)</td>
<td>22.49</td>
</tr>
</tbody>
</table>

D+L Overall: (I-squared = 0.0%, p = 0.982)

M-H Overall: 0.17 (0.06, 0.47)

NOTE: Weights are from random effects analysis
# INH 12-72 vs. RMP-INH 3-4 months

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>OR (95% CI)</th>
<th>(D+L)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Martinez Alfaro</td>
<td>2000</td>
<td>0.30 (0.09, 0.99)</td>
<td>22.06</td>
</tr>
<tr>
<td>Martinson</td>
<td>2011</td>
<td>0.18 (0.09, 0.33)</td>
<td>77.94</td>
</tr>
<tr>
<td>D+L Overall</td>
<td>(I-squared = 0.0%, p = 0.452)</td>
<td>0.20 (0.11, 0.35)</td>
<td>100.00</td>
</tr>
<tr>
<td>M-H Overall</td>
<td></td>
<td>0.20 (0.11, 0.35)</td>
<td></td>
</tr>
</tbody>
</table>

NOTE: Weights are from random effects analysis

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### Graphical Representation

![Graphical representation of INH 12-72 vs. RMP-INH 3-4 months](image-url)