

GRADE Table: Strategies for optimizing HIV monitoring among adults, children and pregnant women living with HIV receiving antiretroviral therapy

| Quality assessment | | | | | | | Summary of findings | Effect | | | | |
|--|---------------|-------------------------|------------------------------|-----------------------------|-------------------------|----------------------|--|---|-------------------|---------------|---------------|------------|
| Question 1.2: Routine CD4 count monitoring versus routine viral load monitoring for adults living with HIV receiving HAART | | | | | | | | | | | | |
| Number of studies | Design | Limitations | Inconsistency | Indirectness | Imprecision | Other considerations | No. of patients CD4 | No. of patients viral load | Relative (95% CI) | Absolute | Quality | Importance |
| Mortality | | | | | | | | | | | | |
| 3 | RCT | Serious limitations (1) | No serious inconsistency | No serious indirectness | No serious imprecision | none | Not available | Not available | Not available | Not available | Moderate-high | Critical |
| New AIDS-defining illness | | | | | | | | | | | | |
| 2 | RCT | Serious limitations (1) | No serious inconsistency | No serious indirectness | No serious imprecision | none | Not available | Not available | Not available | Not available | Moderate | Critical |
| Mortality (observational) | | | | | | | | | | | | |
| 1 | Observational | Serious limitations (2) | No serious inconsistency | No serious indirectness | Serious imprecision (3) | none | Not available | Not available | 0.58 (0.50-0.66) | Not available | Low | Important |
| Clinical monitoring versus routine CD4 count + clinical monitoring | | | | | | | CD4 + clinical monitoring | Clinical monitoring | | | | |
| Switch rates | | | | | | | | | | | | |
| 1 | RCT | No serious limitations | No serious inconsistency | No serious indirectness | Serious imprecision (3) | none | 216/957 (23%) | 173/939 (18%) | 0.75 (0.60-0.96) | Not available | Moderate | Critical |
| Viral suppression (<400 copies/ml) at 5 years | | | | | | | | | | | | |
| 1 | RCT | No serious limitations | No serious inconsistency (4) | No serious indirectness | Serious imprecision (3) | none | 624/737 (85%) | 565/722 (78%) | 0.66 (0.50-0.87) | Not available | Moderate | Critical |
| Viral failure (two consecutive viral loads >500 copies/ml after 90 days) | | | | | | | | | | | | |
| 1 | RCT | No serious limitations | No serious inconsistency (4) | No serious indirectness (5) | Serious imprecision (3) | none | 26/346 (8%) | 19/352 (5%) | Not available | Not available | Moderate | Critical |
| Severe morbidity and mortality | | | | | | | | | | | | |
| 2 | RCT | No serious limitations | Serious inconsistency (6) | No serious indirectness | No serious imprecision | none | 414/2027 (20%) | 531/2033 (26%) | Not available | Not available | Moderate | Critical |
| Routine laboratory (viral load + CD4) monitoring versus clinical monitoring | | | | | | | No. of patients routine laboratory monitoring | No. of patients clinical monitoring | | | | |
| Mortality at 2 years | | | | | | | | | | | | |
| 1 | RCT | No serious limitations | No serious inconsistency | No serious indirectness | Serious imprecision (3) | none | 32/221 (14%) | 44/238 (18%) | 1.31 (0.83-2.06) | Not available | Moderate | Critical |
| Viral suppression (<40 copies/ml) at month 24 | | | | | | | | | | | | |
| 1 | RCT | No serious limitations | No serious inconsistency | No serious indirectness | Serious imprecision (3) | none | 115/221 (52%) | 111/238 (47%) | 0.81 (0.56-1.16) | Not available | Moderate | Critical |
| Mean CD4 cell count increase at 24 months | | | | | | | Mean CD4 cells/mm³ (lab) | Mean CD4 cells/mm³ (clin) | | | | |
| 1 | RCT | No serious limitations | No serious inconsistency | No serious indirectness | Serious imprecision (3) | none | 206 | 175 | -31 (-63-2.0) | Not available | Moderate | Critical |
| Question 1.3: Higher versus lower viral load threshold for switching HAART regimens | | | | | | | | | | | | |

| | | | | | | | No. of patients Lower viral load | No. of patients Higher viral load | | | | |
|---|---|----------------------------|-----------------------------|----------------------------|----------------------------|--|--|---|----------------------|--------------------|----------|----------|
| New CDC Grade 3 or 4 adverse event among children | | | | | | | | | | | | |
| 1 | RCT | No serious limitations | No serious inconsistency | No serious indirectness | Serious imprecision (3) | none | 30/125 (24%) | 30/121 (25%) | 1.05 (0.63- 1.73) | 8 more per 1000 | Moderate | Critical |
| New serious adverse event among children | | | | | | | | | | | | |
| 1 | RCT | No serious limitations | No serious inconsistency | No serious indirectness | Serious imprecision (3) | none | 19/125 (15%) | 29/121 (16%) | Not available | 9 more per 1000 | | |
| New serious adverse event among children (observational) | | | | | | | | | | | | |
| 1 | Observational/secondary modelling analysis | Serious limitations (4) | No serious inconsistency | No serious indirectness | Serious imprecision (3) | Secondary analysis of observational trial (6) | 67 | 67 | Not available | Not available | Low | Critical |

(1) Only Jourdain 2011 reported double-blinding, but did not describe the blinding procedure. In addition, the Jourdain methods have not yet been fully reported, so we downgraded for this.

(2) Keiser 2011 compares the results from three countries in an ecological study; one uses routine viral load monitoring (South Africa), and two do not (Zambia and Malawi).

(3) The results are from one trial only, and therefore the results may be imprecise.

(4) Although Mermin (2011) and Kityo (2012) report viral failure and suppression using different definitions, these findings are consistent with one another.

(5) Although Kityo (unpublished) did not include morbidity or mortality outcomes, given the importance of viral failure in clinical management, we did not downgrade for this.

(6) The study used models of using different viral load cutoffs in a secondary analysis of an observational trial.