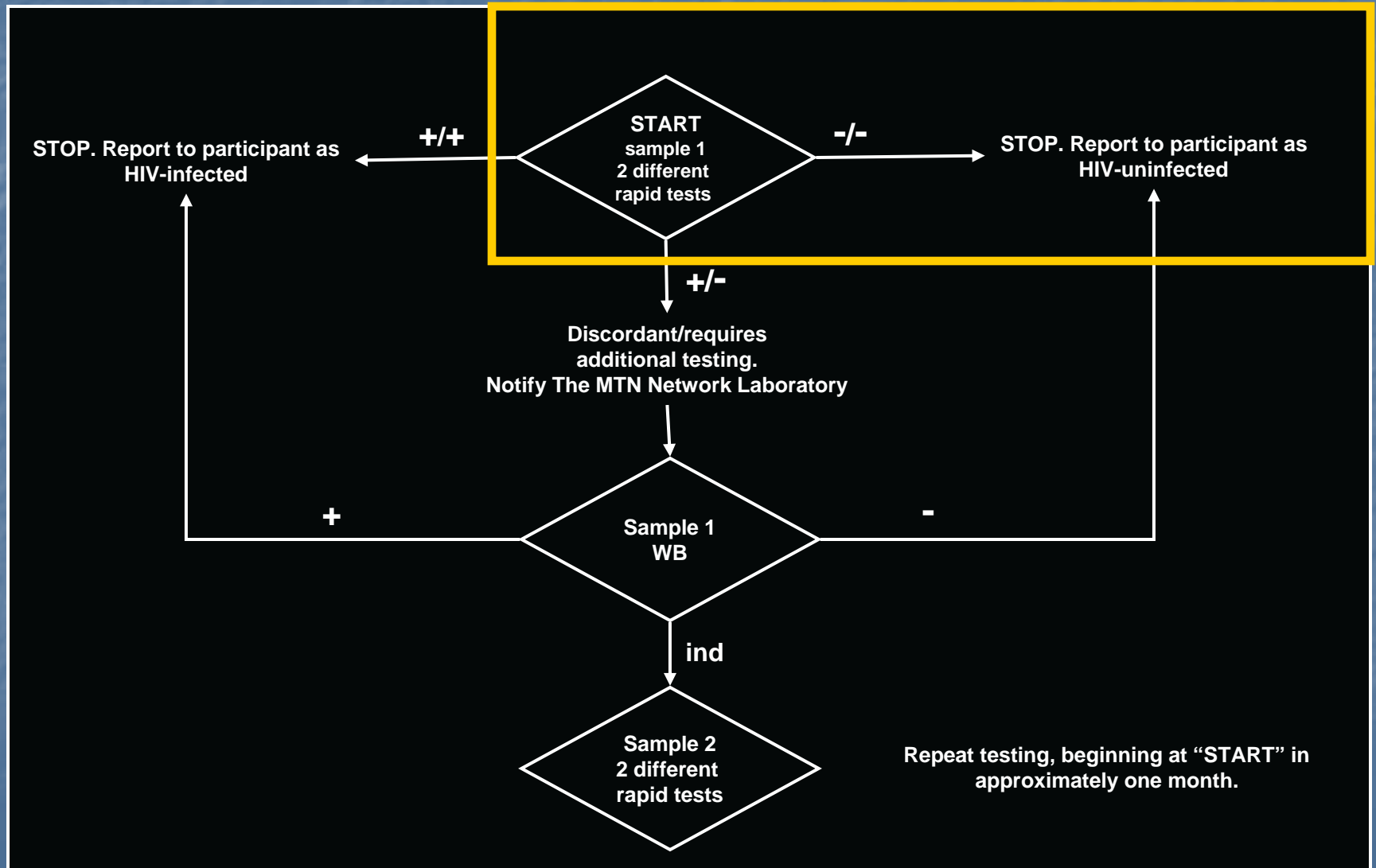


HIV Testing and Study Procedures for Seroconverters

**MTN-003 Operational Walkthrough
11 November 2008
Johannesburg, South Africa**

HIV Testing at Screening

HIV Testing at Screening

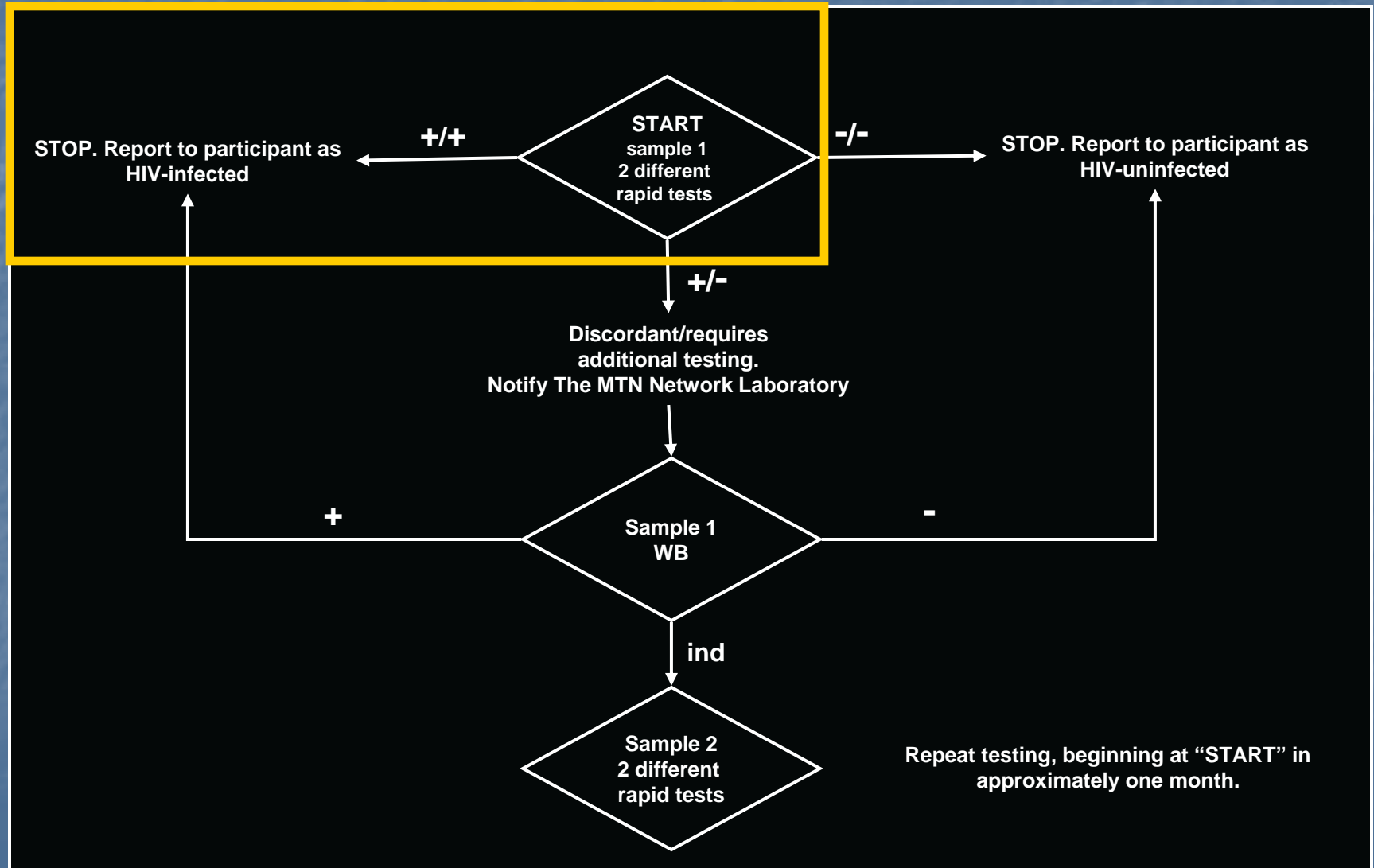


HIV Testing at Screening

Start with two different concurrent rapid tests:

- If both tests **negative** \Rightarrow uninfected \Rightarrow eligible

HIV Testing at Screening

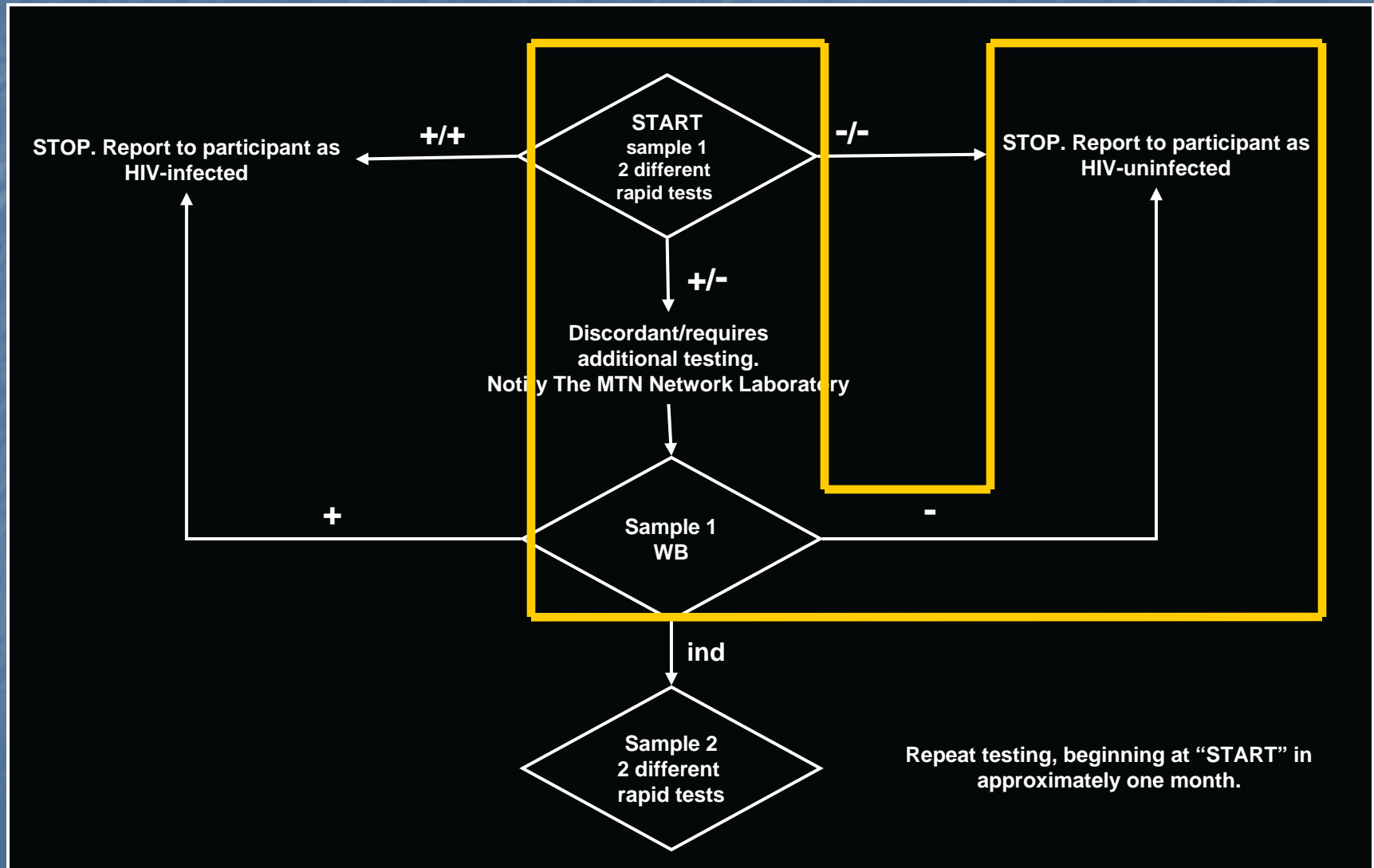


HIV Testing at Screening

Start with two different concurrent rapid tests:

- If both tests **negative** \Rightarrow uninfected \Rightarrow eligible
- If both tests **positive** \Rightarrow infected \Rightarrow not eligible

HIV Testing at Screening

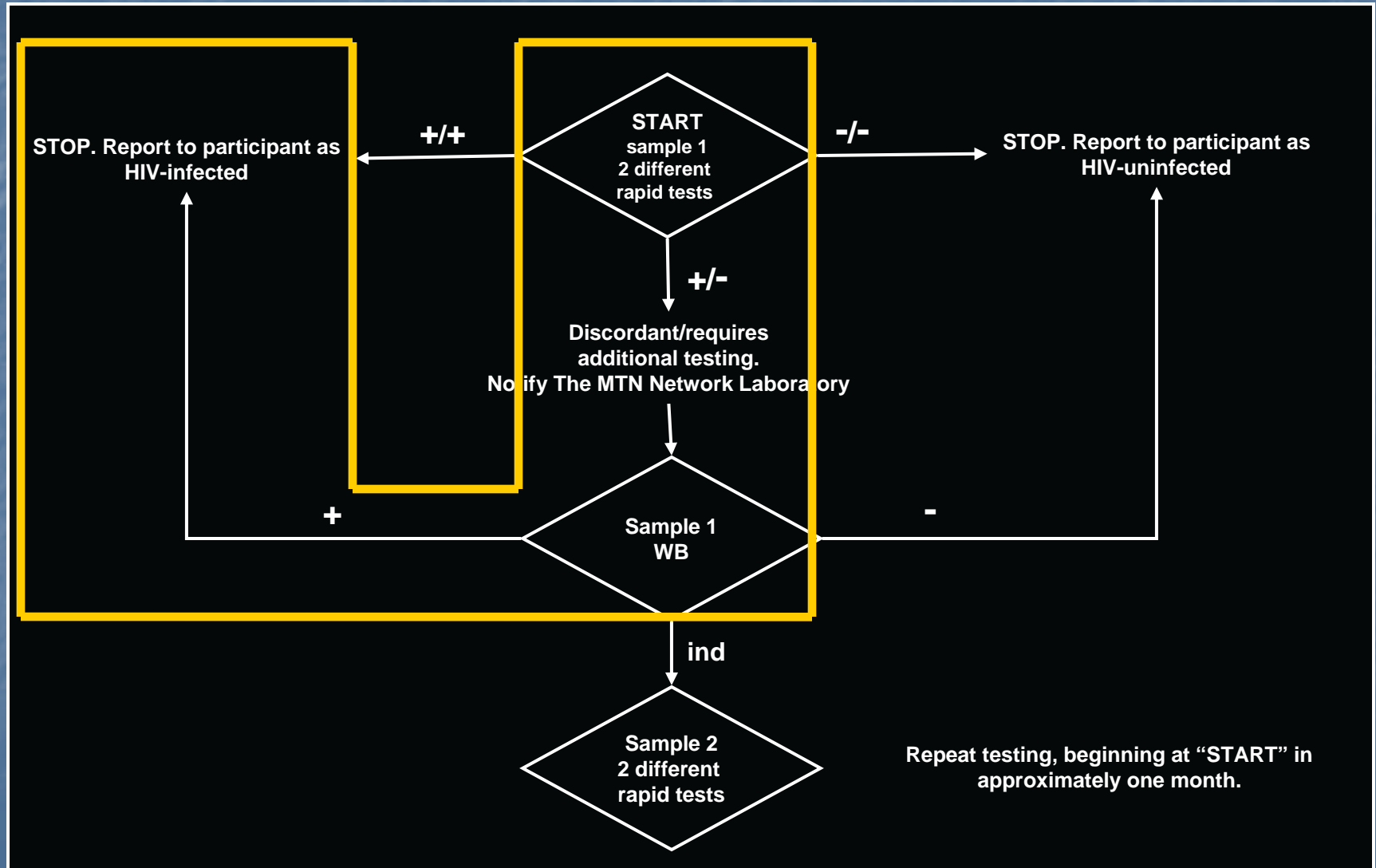


HIV Testing at Screening

Start with two different concurrent rapid tests:

- If both tests **negative** \Rightarrow uninfected \Rightarrow eligible
- If both tests **positive** \Rightarrow infected \Rightarrow not eligible
- If one test **negative** and one test **positive** \Rightarrow WB and notify the MTN NL for their awareness
 - If WB **negative** \Rightarrow uninfected \Rightarrow eligible

HIV Testing at Screening

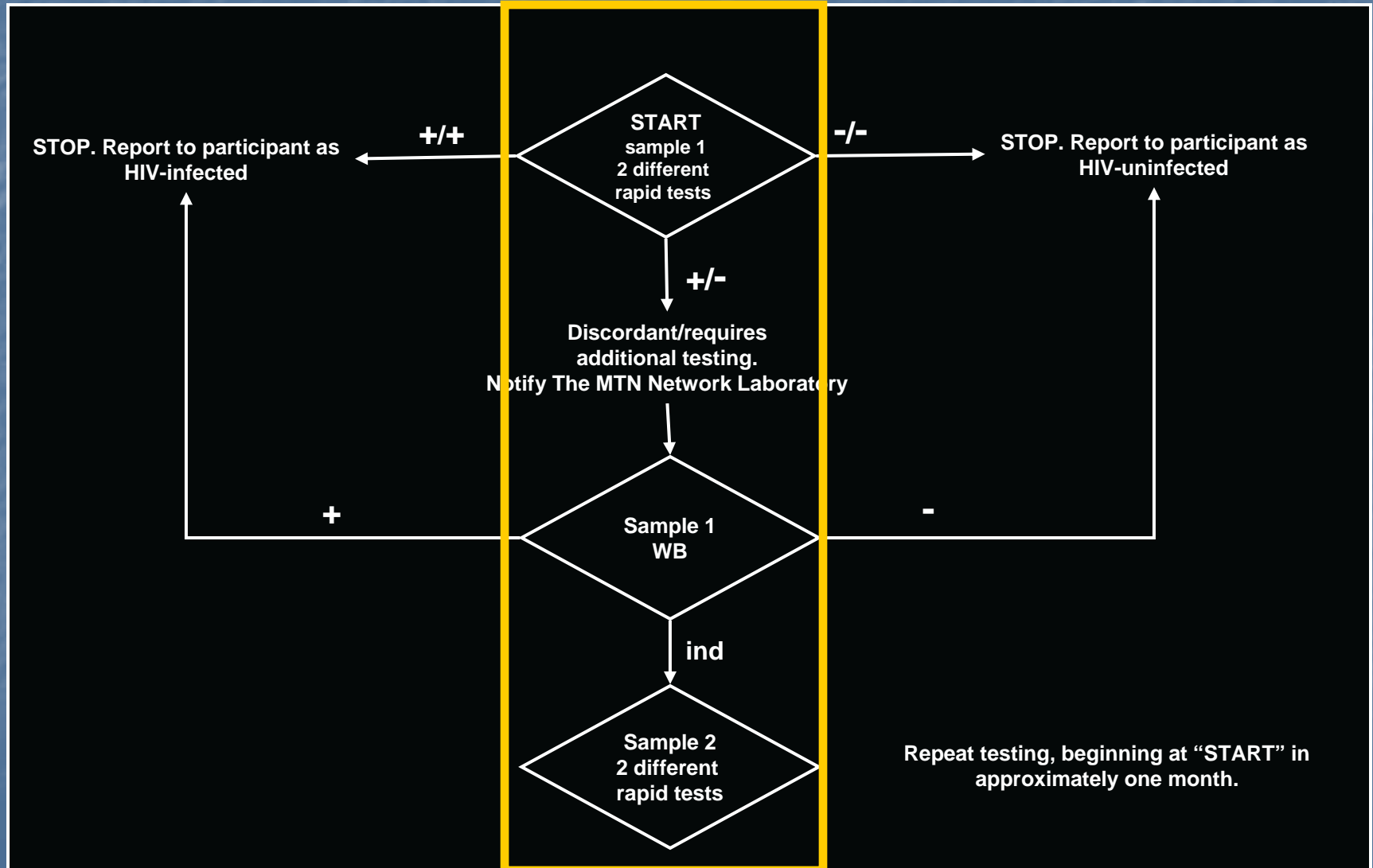


HIV Testing at Screening

Start with two different concurrent rapid tests:

- If both tests **negative** \Rightarrow uninfected \Rightarrow eligible
- If both tests **positive** \Rightarrow infected \Rightarrow not eligible
- If one test **negative** and one test **positive** \Rightarrow WB
 - If WB **negative** \Rightarrow uninfected \Rightarrow eligible
 - If WB **positive** \Rightarrow infected \Rightarrow not eligible

HIV Testing at Screening



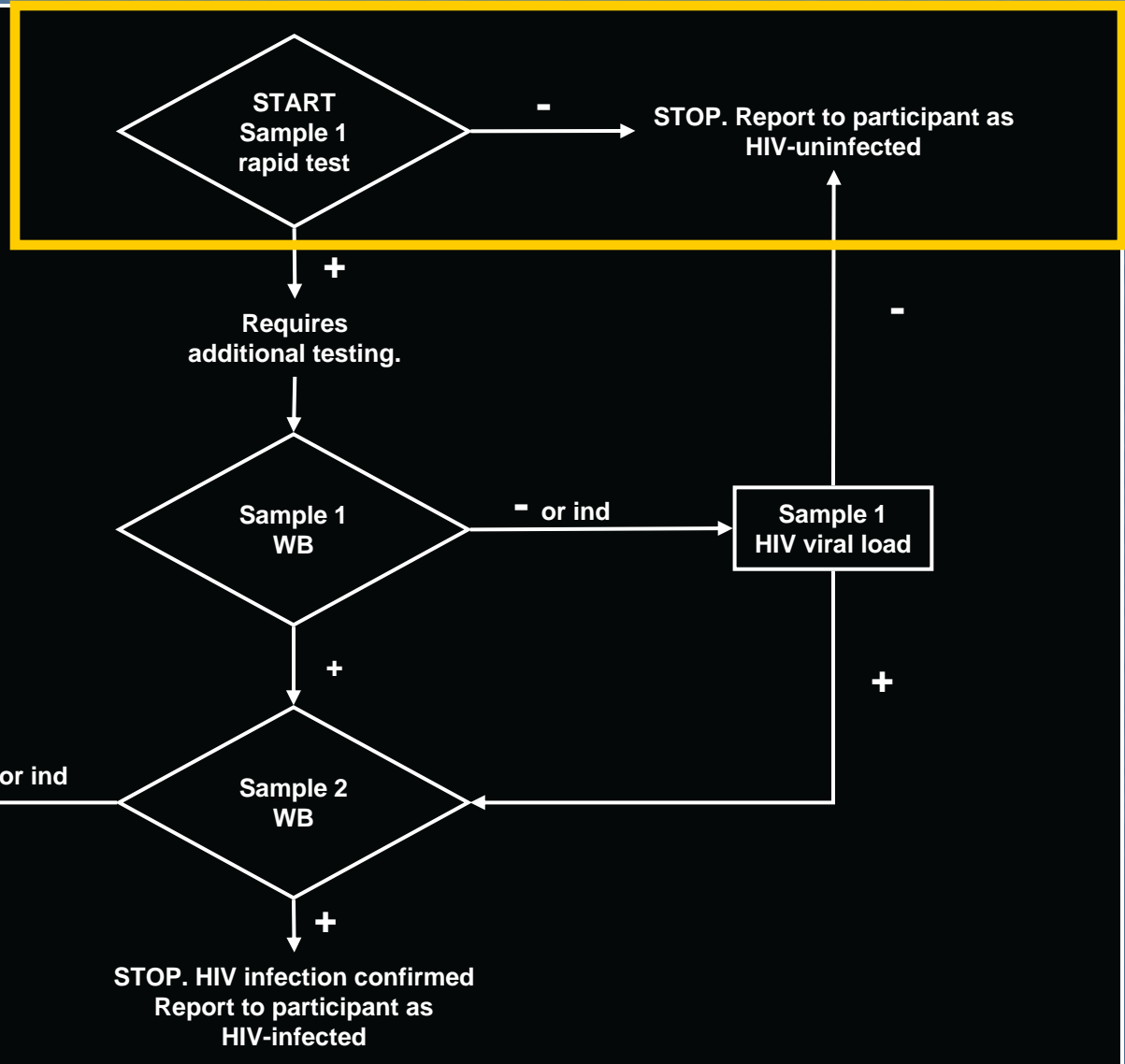
HIV Testing at Screening

Start with two different concurrent rapid tests:

- If both tests **negative** \Rightarrow uninfected \Rightarrow eligible
- If both tests **positive** \Rightarrow infected \Rightarrow not eligible
- If one test **negative** and one test **positive** \Rightarrow WB
 - If WB **negative** \Rightarrow uninfected \Rightarrow eligible
 - If WB **positive** \Rightarrow infected \Rightarrow not eligible
 - If WB **indeterminate** \Rightarrow status unclear \Rightarrow not currently eligible \Rightarrow repeat testing from beginning of algorithm in one month

HIV Testing During Follow-up

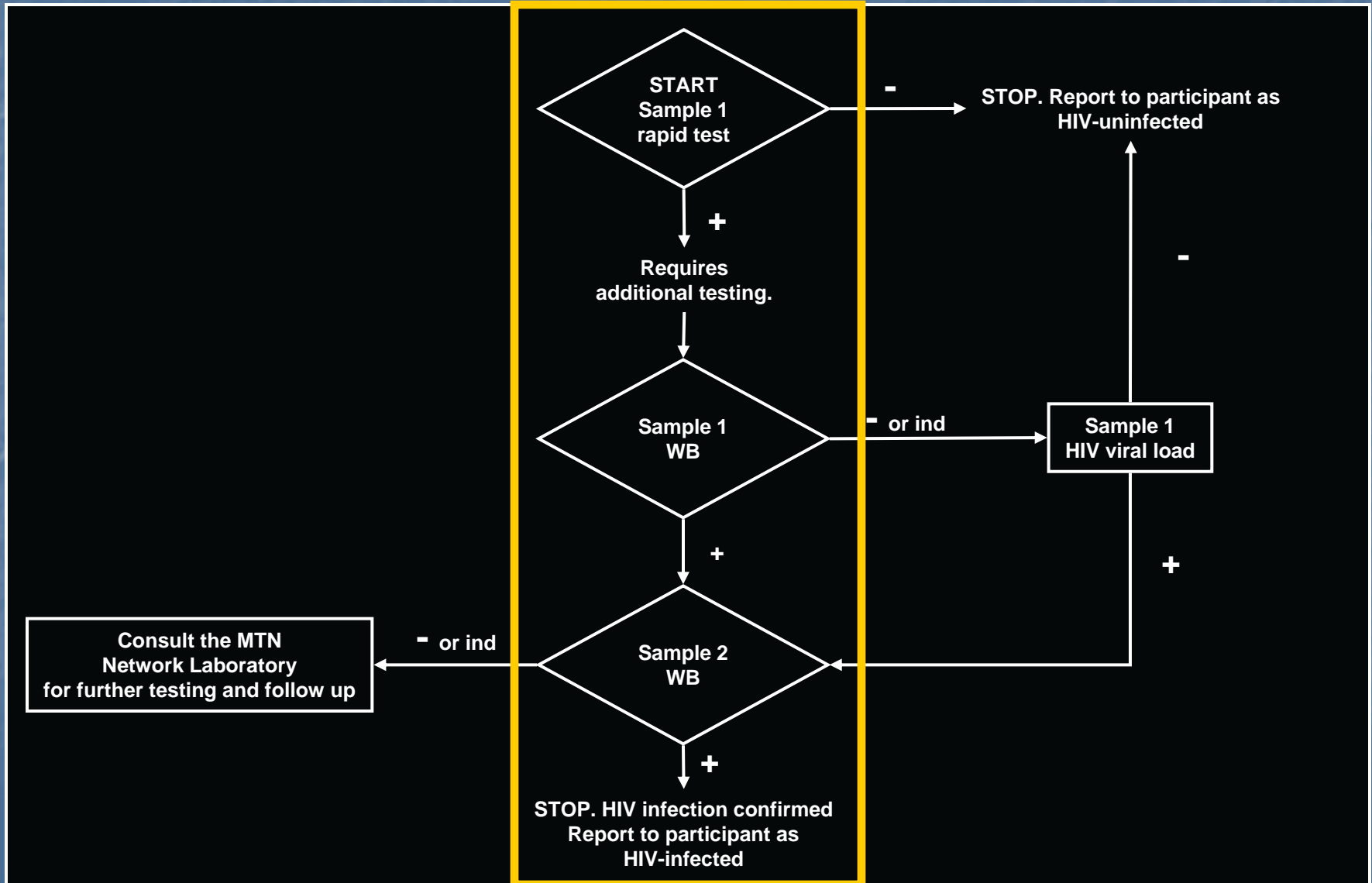
HIV Testing During Follow-Up



HIV Testing During Follow-Up

- Start with rapid test(s)
- If all rapid tests **negative**, report as uninfected, no further testing required

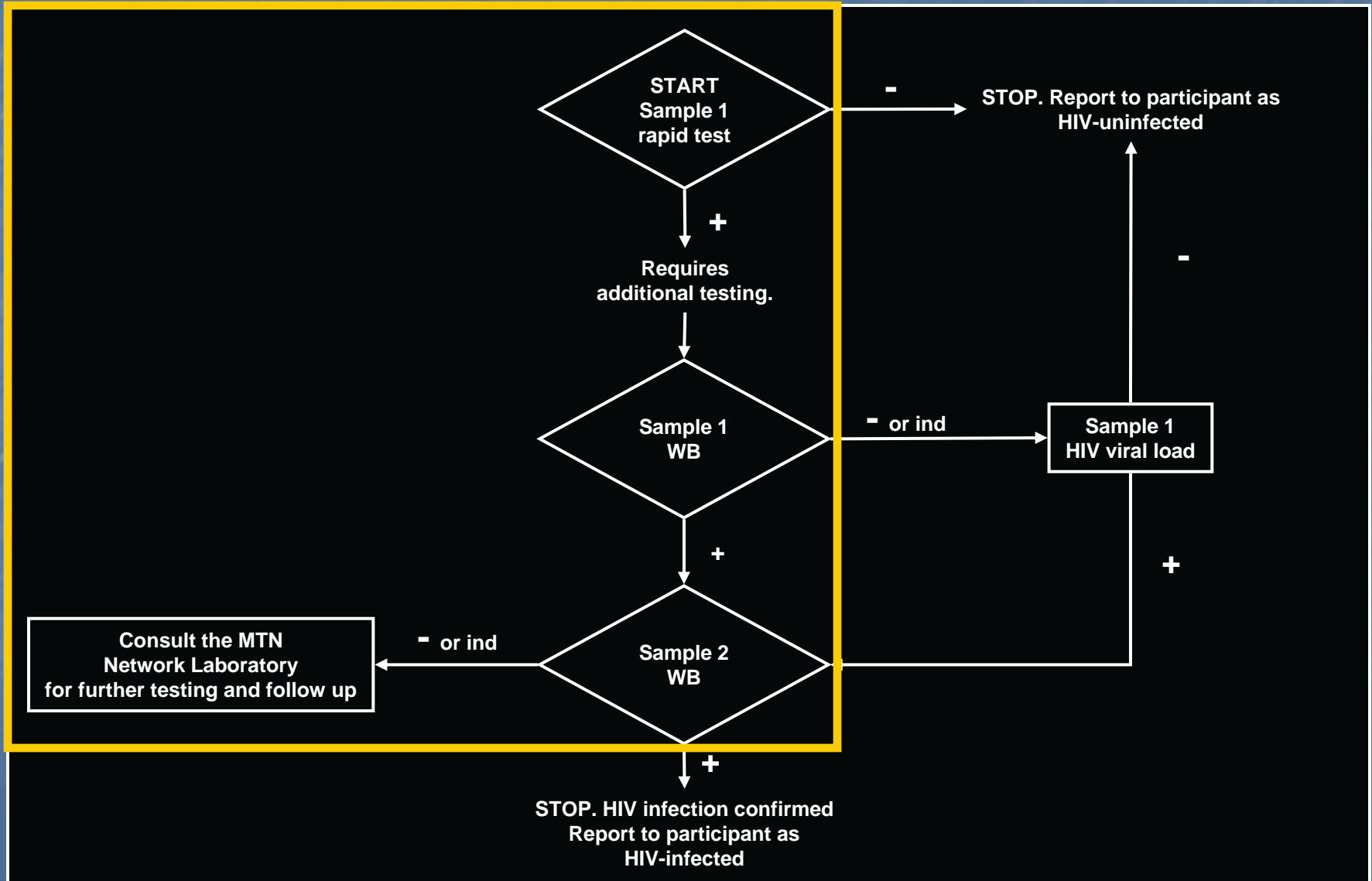
HIV Testing During Follow-Up



HIV Testing During Follow-Up

- Start with rapid test(s)
- If one or more rapid tests **positive**, report as requiring additional testing, request WB (sample 1)
 - If sample 1 WB **positive** \Rightarrow report as infected, collect sample 2 and request WB
 - If sample 2 WB **positive** \Rightarrow seroconversion confirmed, no further testing required

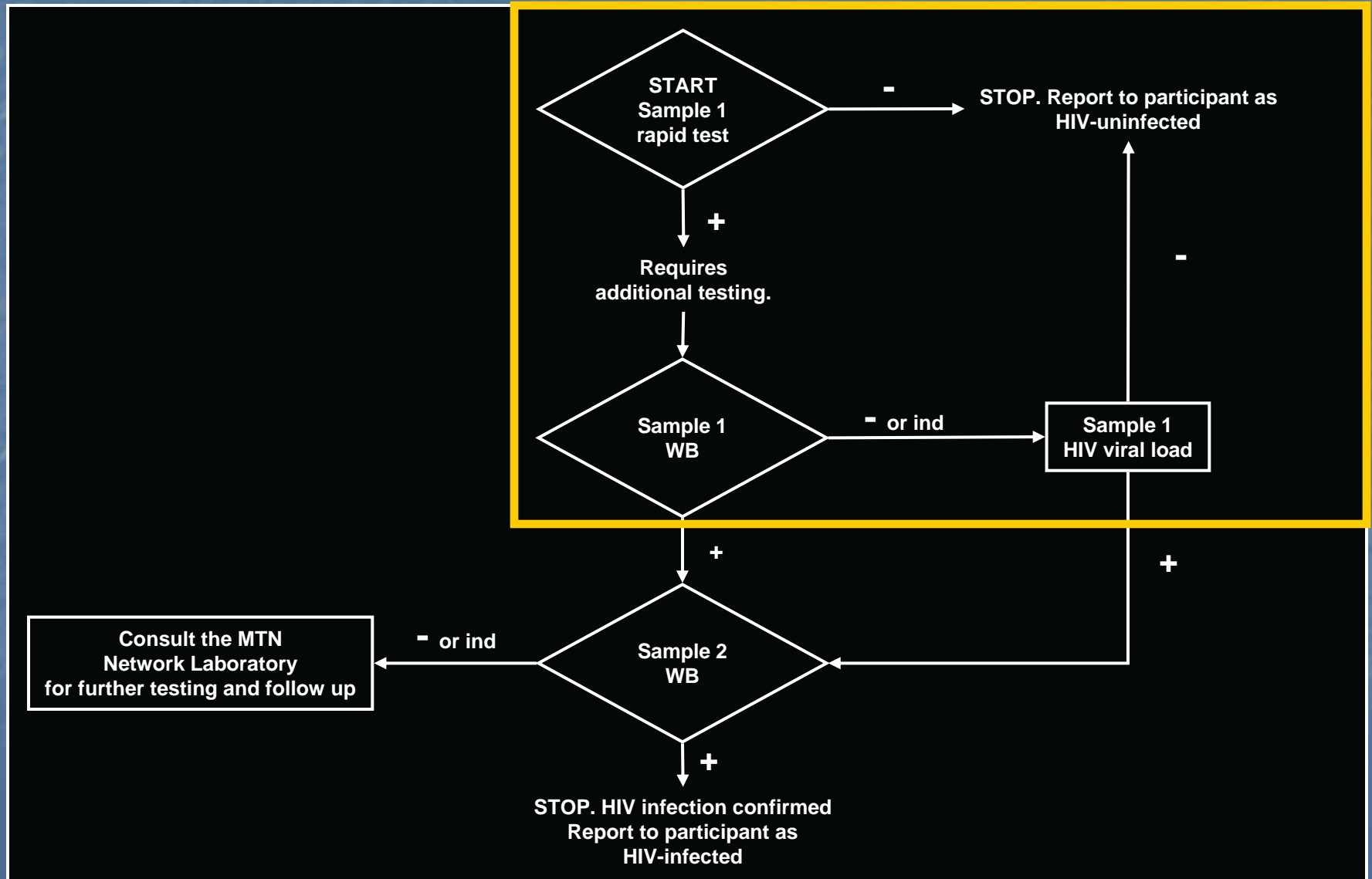
HIV Testing During Follow-Up



HIV Testing During Follow-Up

- Start with rapid test(s)
- If one or more rapid tests **positive**, report as requiring additional testing, request WB (sample 1)
 - If sample 1 WB **positive** ⇒ report as infected, collect sample 2 and request WB
 - If sample 2 WB **negative or determinate** ⇒ status unknown, consult MTN NL

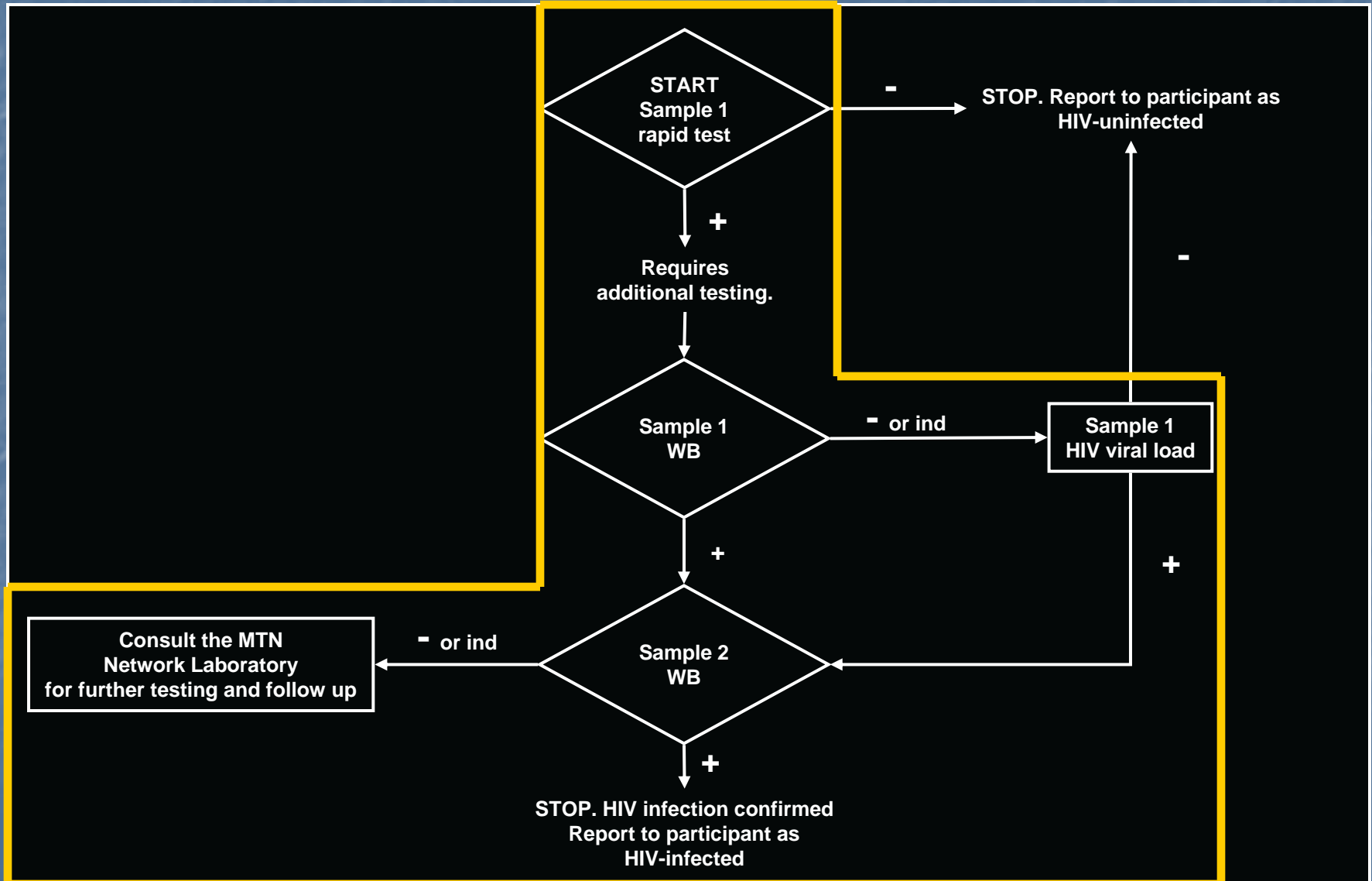
HIV Testing During Follow-Up



HIV Testing During Follow-Up

- Start with rapid test(s)
- If one or more rapid tests **positive**, report as requiring additional testing, request WB (sample 1)
 - If WB **negative** or **indeterminate** ⇒ HIV RNA PCR
 - If PCR **negative**, i.e., result is below the limit of detection for the test, report as uninfected, no further testing required

HIV Testing During Follow-Up



HIV Testing During Follow-Up

- Perform rapid test(s)
- If one or more rapid tests **positive**, report as requiring additional testing, request WB (sample 1)
 - If WB **negative or indeterminate** ⇒ HIV RNA PCR
 - If PCR **positive**, i.e., any detectable amount of HIV, report as requiring additional testing, collect sample 2 and request WB
 - If sample 2 WB **positive** ⇒ seroconversion confirmed, no further testing required
 - If sample 2 WB **indeterminate** ⇒ status unknown, consult MTN NL

Procedures for (Possible) Seroconverters

*** currently under review ***

- Per protocol Section 7.5.3, additional procedures must be performed while seroconversion is being confirmed:
 - CD4 cell count
 - HIV RNA PCR
 - Plasma archive for confirmatory HIV testing at the MTN NL **and for resistance testing at the MTN NL**
 - ~~■ PBMC archive~~
 - **All done when blood is collected [each sample] for Sample 2**

Sample 2 Tips and Reminders

- Collect sample 2 at visit when participant receives her sample 1 WB and PCR results
- Collect approximately **15 mL** of whole blood
*** currently under review ***
- In the lab, aliquot blood needed for WB, HIV RNA PCR, and CD4 cell count from one tube
- Then process all remaining blood for plasma archive

Procedures for Seroconverters

- Per protocol Section 7.6.1, seroconverters will be offered the option to continue follow-up until their originally scheduled study exit date
- For those who continue follow-up, all procedures continue except:
 - HIV serology
 - Provision of study product and other product-related procedures
- HIV/STI risk reduction counseling will be modified to address primary and secondary prevention for infected women

Procedures for Seroconverters

*** currently under review ***

- Per protocol Section 7.6.1, seroconverters will be offered enrollment in MTN-015
- Participants who delay or decline enrollment in MTN-015 undergo the following as part of VOICE:
 - CD4 cell count, HIV RNA PCR, plasma archive, ~~PBMC archive~~
 - At Months 1, 3, 6, and q6 months post-seroconversion
 - Plasma and ~~PBMCs~~ will be shipped to the MTN NL for resistance testing, testing of TDF and FTC levels, and long-term archive and possible future research testing

Procedures for Seroconverters

- For the above-listed “post-seroconversion” procedures, the date of seroconversion is the specimen collection date of Sample 1
 - **This sets a new scheduling calendar for these procedures**
 - **But all other VOICE visits and procedures remain on the VOICE monthly visit schedule**
- If Month 1 post-seroconversion has passed by the time seroconversion is confirmed, the Month 1 procedures may be omitted

**What Are
Your Questions?**