# NMDP HML Gateway Programmatic Validation

## 09-April-2020

HML Gateway's /validate endpoint performs schema, internal schema, and semantic validation. Incoming HML undergoes an initial sanity check, a structure/schema validation, and a rules/semantics validation. Internal schema validation ascertains gene family consistency.

#### **Sanity Check**

- The submitted HML cannot be null or empty
- The submitted HML's HML ID cannot be null
- The submitted HML's HML ID root value cannot be null or empty
- The submitted HML must have at least one sample
- The submitted HML version must be the currently supported version, 1.0.1

## **Schema Validation**

This isn't simply binding the incoming HML to a java object, HML gateway will produce a list of schema validation errors. This list can be used without having to read the schema file manually to figure out where the HML is malformed.

## **Internal Schema Validation**

The HML's 'gene-family' typing element attributes must all have the same value

## **Semantic Validation**

- The HML must be correctly bound to HML Gateway's Java class. The Java HML object cannot be null
- The HML reporting center value cannot be null or empty
- The HML project name must be valid (see section on project validation)
- The HML version cannot be null or empty
- The HML ID cannot be null or empty
- The HML must have no duplicate sample IDs
- An HML sample's center code must be lower or uppercase alpha, or numeric, and have no more than 50 characters
- All HML submission samples must have the same gene family
- Each reference sequence start position cannot be greater than the reference sequence end position
- Each variant start position cannot be greater than the variant end position
- Each HML sample must have at least one typing
- Each typing date cannot be null
- Each typing date cannot be set in the future of the submission time
- Each typing method cannot be empty or null

- Each typing method must be either SSO, SSP, SBT-Sanger, or SBT-NGS (i.e. must be one of the following lowercase values: "sso", "ssp", "sbt-sanger", "sbt-ngs")
- If the typing method test-id is present, test-id-source must be also
- If the typing method test-id-source is present, test-id must be also
- Finally, each HML instance must pass additional semantic rule validation for HLA, or KIR

## **Project Name Semantic Validation**

- The HML submission project name cannot be null or empty
- For NMDP Form 22 submissions, the project name must be "REG CONFIRM DNR"
- For NMDP Form 117 submissions, the project name must be "REG CONFIRM RCP"
- For CIBMTR form 2005 submission, project name is a research project name, i.e. it has a "RES\_" prefix and is comprised of only lower and uppercase letters, or numbers (0-9), or the "-" character. (see separate instructions for CIBMTR Form 2005 submissions).

#### **HLA Semantic Validation**

- An HML sample requires at least one allele assignment
- An HML sample's allele assignment date cannot be null
- An HML sample's allele assignment date cannot be set in the future of the submission time
- An HML sample's haploid type cannot be equal to "0000". Will be reported as "FAILED-RESULT"
- An HML sample's GL String cannot represent more than one HLA locus
- An HML sample's locus cannot be reported more than once
- An HML sample's GL String cannot contain both a text value and a URI
- A GL String must be Uppercase alpha, numeric, or one of the following characters: +, ^,
  ~, |, \*, :, -, (or a character that does not match the \w regex character)
- A GL String must contain at least one fully qualified allele name (i.e. this GL String must contain at least one "\*" character)
- A GL String cannot start with a +, /, |,  $\sim$ , or  $^{\land}$  character
- A GL String cannot end with a +, /, |,  $\sim$ , or  $^{\land}$  character
- A GL String allele name must match the following regex: "(HLA\-)(A|B|C|DRB1|DRB3|DRB4|DRB5|DQB1|DPB1|DQA1|DPA1)\s?(\d{2,5})(:\d{2,5})(:\d{2,5})(+"\d{2,5})(-\d{2,5})
- An allele name cannot be less than one character
- Genotype lists are not supported, so they are ignored
- An allele assignment must have at least one haploid or GL string
- A non-even number of haploids will generate a warning
- An HML sample's haploid type cannot be blank
- HML sample haploids cannot have more than two alleles for the same locus even across multiple typings
- A GL String cannot have more than two alleles for the same locus (i.e. for any substring surrounded by | characters, no more than one + or ~ character delimiting 2 allele substrings)

## **KIR Semantic Validation**

- Allele assignment haploids are not allowed for gene-family KIR
- A KIR GL String cannot contain both a text value and a URI
- Typing allele assignment properties must have valid supply order line IDs