Clinical suspicion or family history of Fabry disease

- **FMTT / Familial Mutation, Targeted Testing, Varies**
  - **YES**
    - Positive family history with mutation identified
      - Males
        - Normal alpha-galactosidase enzyme activity
          - Fabry disease highly unlikely (<1% of affected males have leukocyte enzyme values within the normal range)
        - Decreased alpha-galactosidase enzyme activity
          - Consider: FMTT for at-risk family members
      - Females
        - Analysis of alpha-galactosidase enzyme activity is not sensitive for carrier detection in all females. It is recommended that molecular testing be performed for diagnosis in females.
            - No variant or pseudodeficiency allele identified
              - Fabry disease unlikely. Results must be interpreted in the context of an individual's clinical and/or biochemical profile.
            - Disease-causing variant identified
              - Fabry disease confirmed
            - Variant of unknown significance identified
              - If not already performed, consider: LGB3S / Globotriaosylsphingosine, Serum
                - No variant identified
                  - No variant or pseudodeficiency allele identified
                    - Fabry disease unlikely. Results must be interpreted in the context of an individual's clinical and/or biochemical profile.
                  - Disease-causing variant identified
                    - Fabry disease confirmed
                  - Variant of unknown significance identified
                    - If not already performed, consider: LGB3S / Globotriaosylsphingosine, Serum
                      - No variant identified
            - Variant of unknown significance identified
              - If not already performed, consider: LGB3S / Globotriaosylsphingosine, Serum
                - No variant identified

- **NO**
  - Recommended initial evaluation (select 1)
    - AGAW / Alpha-Galactosidase, Leukocytes and/or
    - AGAS / Alpha-Galactosidase, Serum
  - Normal alpha-galactosidase enzyme activity
  - Decreased alpha-galactosidase enzyme activity

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