VACCINE-INDUCED PROTHROMBOTIC IMMUNE THROMBOCYTOPENIA (VIPIT)



OBJECTIVE:

To assist health care professionals in the diagnosis and management of Vaccine-Induced Prothrombotic Immune Thrombocytopenia (VIPIT) also known as Vaccine-induced Immune Thrombotic Thrombocytopenia (VITT).

BACKGROUND:

Vaccines are a critical tool in the management of the COVID-19 pandemic resulting from SARS-CoV-2. Several vaccines have been rapidly developed and subsequently approved by Health Canada and deployed across Canada. Among those is the ChAdOx1 nCoV-19 vaccine (AZD1222) developed at Oxford University and produced by AstraZeneca and the Serum Institute of India.

Recently, after widespread vaccination with the AstraZeneca vaccine in Europe, there have been reports of some vaccine recipients developing unusual thrombotic events and thrombocytopenia. Investigators have concluded that the AstraZeneca vaccine is associated with development of a prothrombotic disorder that clinically resembles heparin-induced thrombocytopenia (HIT). [See Clinical Guide Heparin Induced Thrombocytopenia.]

DIAGNOSIS OF VACCINE-INDUCED PROTHROMBOTIC IMMUNE THROMBOCYTOPENIA (VIPIT)/ VACCINE-INDUCED IMMUNE THROMBOTIC THROMBOCYTOPENIA (VITT):

Patients presenting with the following blood clotting symptoms should be asked about their vaccine history:

- a persistent and severe headache
- focal neurological symptoms or visual changes, including blurred or double vision, or episodes suspicious for seizure
- shortness of breath
- abdominal or chest pain
- swelling and redness in a limb
- pallor and coldness in a limb
- unusual bleeding, multiple small bruises, reddish or purplish spots or blood blisters under the skin

If their vaccination falls within the period of 4-28 days prior to presentation, a complete blood count (CBC) should be drawn. If the platelet count is $< 150 \times 10^9 / L$ and the vaccination window is between 4 and 28 days, the patient should be evaluated at their nearest emergency department for suspected VIPIT/VITT. Patients whose vaccination was not within the 4-28 day period or whose platelet count is $> 150 \times 10^9 / L$ are unlikely to have VIPIT/VITT.

Patients with suspected VIPIT/VITT should go on to have a D-dimer level and a blood film. In cases where there is strong clinical suspicion of VIPIT/VITT, patients should also have diagnostic imaging to investigate for blood clots (including appropriate imaging to rule out cerebral vein sinus thrombosis

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(CVST), if the patient presents with a persistent and severe headache, nausea/vomiting, visual changes, focal neurological deficits or episodes suspicious for seizures. It is prudent to note that imaging to rule out CVST includes both parenchymal imaging and vascular imaging, either with a CT head/CT venogram, or MR head/MR venogram and that this potential diagnosis should be investigated urgently with same-day neuroimaging. It is not known whether VIPIT/VITT, like HIT, is associated with arterial thromboses, but arterial clots should be considered if patients have consistent symptoms. Low platelet count, abnormal coagulation parameters, a normal blood film (apart from thrombocytopenia), with or without confirmation of a blood clot on diagnostic imaging makes the diagnosis of VIPIT/VITT presumptive. The confirmatory diagnosis of VIPIT/VITT is made by testing for HIT. [See Clinical Guide Heparin Induced Thrombocytopenia].

Do not proceed to HIT testing No Work up alternate Symptoms of VIPIT/VITT diagnoses persistent and severe headache; focal neurological symptoms or Onset between visual changes, including blurred 4 and 28 days or double vision, or episodes after suspicious for seizure; shortness of vaccination breath; abdominal or chest pain; swelling and redness in a limb; or pallor and coldness in a limb **Laboratory Investigations:** Yes Platelet Count D-Dimer Low platelet count <150 x 10⁹/L D-dimer: elevated Blood film: normal (apart from low platelet) Imaging based on clinical suspicion confirms venous or arterial No thrombosis ** Not all cases of VITT initially present with a clot. A confirmed blood clot is not required to make a presumptive diagnosis of VITT ** Presumptive diagnosis of VIPIT/VITT Consult hematology and proceed to HIT **ELISA** testing Recommended Laboratory Methods: 1. Antigen-binding assay (ELISA) for Yes PF4/heparin antibodies: ELISA Testing 2. NOTE: Rapid immunoassay (RSA), chemiluminescence immunoassay (CLIA), may reveal false-negative Send for testing before initiation of treatment. Do not wait for test results before initiation of

Figure 1: Decision Tree for Diagnosing and Ruling Out VIPIT/VITT

Image adapted and used with permission from Ontario's COVID-19 Science Advisory Table

MANAGEMENT OF VIPIT/VITT

Given the uncommon nature of VIPIT/VITT, at the time of presumptive diagnosis, clinicians should refer for an urgent hematology consultation. Patients with presumptive and confirmed VIPIT/VITT should be treated similarly to HIT.

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Treating Patients with Presumptive or Confirmed VIPIT/VITT

- No heparin
- No platelet transfusions
- First line anticoagulants: direct oral factor Xa inhibitors (e.g., rivaroxaban, apixaban, edoxaban)
- Consult hematologist
- IVIG 1g/kg daily for at least 2 days

[See also Clinical Guide Heparin Induced Thrombocytopenia].

NOTE: All suspected adverse events following immunization (AEFI), including thrombosis, and both presumptive and confirmed VIPIT/VITT, should be reported using the provincial AEFI form and sent to the local Public Health Unit.

OTHER RELEVANT THROMBOSIS CANADA CLINICAL GUIDES AND RESOURCES:

Clinical Guides:

- Cerebral Venous Thrombosis
- Heparin Induced Thrombocytopenia

Resources:

COVID-19 Vaccines and Blood Clots FAQs

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Please note that the information contained herein is not to be interpreted as an alternative to medical advice from your doctor or other professional healthcare provider. If you have any specific questions about any medical matter, you should consult your doctor or other professional healthcare providers, and as such you should never delay seeking medical advice, disregard medical advice or discontinue medical treatment because of the information contained herein.

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