

Annex 1

Updated (Aug 2021) COVID-19 AESI including status of associated Brighton case definitions

AESI ^{Rationale to include as AESI (1, 2, 3, 4 and/or 5)}	Brighton Case Definition Status
AESI included because they are seen with COVID-19 Disease ^{3,4}	
Acute respiratory distress syndrome	Published ⁶
Multisystem inflammatory syndrome (children & adults)	Published ⁶
1. Myocarditis / pericarditis 2. Other forms of acute cardiac injury including arrhythmias, heart failure, coronary artery disease, myocardial infarction, stress cardiomyopathy	1. Completed, submitted to Vaccine ⁶ 2. No Brighton case definition to be developed but companion guide to myocarditis/pericarditis will include background rates and ICD/MedDRA codes
Coagulation disorder (includes thrombotic disorders, bleeding disorders)	Thrombosis near completion. Bleeding disorder WG to be formed
Anosmia, ageusia	WG formed & developing case definition
Chilblain – like lesions	WG to be formed
Erythema multiforme	Not yet started
Single Organ Cutaneous Vasculitis	Published
Acute kidney injury	Published lab-based criteria (see *)
Acute liver injury	Published lab-based criteria (see #)
Acute pancreatitis	Not yet started
Rhabdomyolysis	Not yet started
Subacute thyroiditis	Not yet started
AESI included because they have a proven or theoretical association with immunization in general	
Anaphylaxis ^{1,2}	Published ⁶
Thrombocytopenia ^{1,2,3,4}	Published ⁶
Generalized convulsion ^{1,2}	Published ⁶
Acute disseminated encephalomyelitis ⁴	Published ⁶
Guillain Barré Syndrome ^{3,4}	Published ⁶
AESI included because they have a proven or theoretical association with specific vaccine platform(s)	
Acute aseptic arthritis ^{r-VSV}	Published ⁶
Aseptic meningitis ^{Live vaccines}	Published ⁶
Encephalitis / Encephalomyelitis ^{Live vaccines}	Published ⁶
Bell's Palsy ^{Intranasal EColi Heat Labile Toxin Adjuvanted Vaccine}	Published ⁶
Vaccine associated enhanced disease ^{1(Formalin inactivated measles/RSV; HIV), 2(Chimeric YF Dengue), 5 (SARS / MERS-CoVs)}	Published ⁶

¹ Proven association with immunization encompassing several different vaccines

² Proven association with vaccine that could theoretically be true for novel COVID-19 vaccines

³ Theoretical concern based on wild type disease immunopathogenesis

⁴ Theoretical concern related to viral replication during wild type disease

⁵ Theoretical concern because it has been demonstrated in an animal model with ≥ 1 vaccine platform

* Acute kidney injury—consensus definition of Kidney Disease Improving Global Outcomes expert consensus group www.kdigo.org

• Increase in serum creatinine by ≥ 0.3 mg/dl (≥26.5 umol/l) within 48 hours; OR

- Increase in serum creatinine to ≥ 1.5 times baseline, known or presumed to have occurred within prior 7 days OR
 - Urine volume ≤ 0.5 ml/ kg/ hour for 6 hours
- # Acute liver injury – definition as used in majority of COVID-19 publications (but no international consensus):
- > 3-fold elevation above the upper normal limit for ALT or AST OR
 - > 2-fold elevation above the upper normal limit for total serum bilirubin or GGT or ALP

⁶Case Definition and resources available at:
https://docs.google.com/spreadsheets/d/1QgF35nYcsaFN3DZT0tV_IP0TYqQzsDMUQBAd5M9brM/edit#gid=1666959512