Guidance Proposal from SSC Malignancy and Haemostasis Subcommittee

"Prevention and treatment of venous thromboembolism in adults receiving asparaginase therapy"

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Background:

- Administration of asparaginase for the treatment of acute lymphoblastic leukemia is associated with high rates of venous thromboembolism complications
 - >30% in analyses of DFCI protocol (Grace et al. BJH 2011)
 - o High rate of cerebral sinus venous thrombosis
- Asparaginase lowers levels of endogenous anticoagulants (antithrombin, protein C and S)
- Number of potential risk factors have been identified including number of doses, age, coadministration of steroids or anthracycline-based chemotherapy
- Approaches to thromboprophylaxis vary considerably across centers
 - Prophylactic strategies evaluated include prophylactic low molecular weight heparin and/or antithrombin concentrate
- Treatment and prevention of asparaginase-induced VTE not addressed in societal guidelines

Guidance document will address the following issues:

- Primary prophylaxis
 - Anticoagulation including which target populations and indications
 - o Laboratory monitoring and repletion of antithrombin, fibrinogen, and platelets
 - Differential approaches for induction or consolidation
- Treatment of asparaginase-related VTE
 - Selection of anticoagulant
 - o Role for antithrombin concentrate and monitoring levels
 - Management of central venous catheter
 - Implications for ongoing use of asparaginase
 - o Duration of anticoagulation