Abstract – This case report discusses an expanding demographic: Aviation-related venothromboembolism (VTE) in pregnancy. As the number of commercial flights increase and the notion of travel gains popularity, the obstetrician is certain to meet an increasing number of at-risk mothers bound for a long haul flight. The pregnancy state itself being hypercoaguable, studies show that air-travel specific conditions namely immobility, dehydration and hypobaria which leads to thrombus formation. Despite VTE being a leading cause of maternal mortality in world wide, specific preventive protocols regarding the at-risk pregnant flier remain scarce-to-nil. Data has shown that compressive stockings alone remain insufficient. The much-debated role of injectable thromboprophylaxis pre-flight is discussed here. Data and studies regarding oral anticoagulants for prevention and treatment of VTE are fast rising and their roles in pregnancy explored. In summation, evidence irrevocably shows that the risks of VTE outweigh that of bleeding. However, the imperative for thromboprophylaxis in at risk pregnant fliers is still controversial. Copyright © 2016 Penerbit Akademia Baru - All rights reserved.

Keywords: veno-thromboembolism, pulmonary embolism, air travel, thromboprophylaxis

1.0 INTRODUCTION

Venothromboembolic diseases remain one of the leading cause for maternal mortality in Malaysia. The hypercoagualable state during pregnancy doubles the risk of thromboembolism when coupled with air travel. A long journey over four hours alongside immobilization and low cabin pressure are the main triggers for thrombus formation. Most aviation companies have yet to produce a standard preventive protocol for the at risk pregnant women taking long haul flights.

2.0 CASE REPORT

A 36 years old Chinese woman in her sixth pregnancy boarded a long haul flight (about 18 hours) in early pregnancy. She was about 13 weeks pregnant at the time, sans prior veno-thromboembolic (VTE) event. History drew nil for calf tenderness, contraceptive usage or surgery before this presentation. Five days after the flight, she presented to the emergency unit
complaining of shortness of breath, which started during the flight and worsening over the past 3 days. Her risk factors then were advanced maternal age, multiparity, obesity, pregnancy and long flight.

![Figure 1: CTPA with arrows showing filling defects bilaterally.](image1)

![Figure 2: CTPA with arrow showing filling defects and thrombus of the left pulmonary artery.](image2)
On admission, she was comfortable in recumbence, comprehensible and speaking with full sentences during the interview. There was no evidence of calf tenderness or positive cardiovascular findings other than tachycardia. The chest x-ray showed cardiomegaly, the lower limbs Doppler ultrasonography was negative for evidence of deep vein thrombosis; echocardiography yielded normal findings. In view of her multiple risk factors for VTE, a Computerized tomography scan of the pulmonary arteries (CTPA) was done and showed extensive bilateral pulmonary embolism involving the main pulmonary arteries extending to its branches (Figure 1, 2, 3).

The patient was started on subcutaneous tinzaparin and under close monitoring by obstetrician and physician throughout her pregnancy.

3.0 DISCUSSION

Pregnancy is known to increase the risk for thromboembolism by 4-10 times [1, 2, 4] compared to the non-pregnant state. This risk is present from the first trimester [3] and is at the highest in the third trimester [2]. Post partum, this risk increases 100-fold in the immediate first week post partum [1] and remains at 20- to 80-fold until 6 weeks post partum [1, 2].

Air travel is one significant factor for VTE risks. Symptomatic VTE in a healthy young individual after a 4-hour flight is 1:6000 [5] and this risk increases with flight duration: 4-folds over 8 hours, and 8-folds over 12 hours [6]. A recent meta-analysis has shown dose relation where the relative risk for thrombosis increases 26% every 2 hours of flight duration [7]. A cohort study [9] estimates that compared to not flying, the risk of thrombosis after long-haul flights remained 3.2-fold increased which normalized over an eight-week window.
Together, pregnancy-related physiological changes and air-travel beyond 4 hours heightens the thromboembolic risk exponentially. It is not solely the immobility of long haul flights that contribute, but also the hypoxic-hypobaric cabin environment [8] as reported by several studies. The risk of venous thrombosis in a pregnant woman travelling by air is calculated to be 1 per 1000 pregnant air-travellers. (The risk of venous thrombosis in non-pill using women who travel by air is 1:5000 travellers [9], while pregnancy leads to increased risk of thrombosis 5-fold [2]).

Hence, this patient was at risk of thromboembolism in view of her multiple risks factors (high BMI, age more than 35, multiparity, long distance travel and ex-smoker). According to Green top guidelines [18], she requires antenatal thromboprophylaxis throughout her pregnancy and postpartum period. However, the role of thromboprophylaxis and mechanical prophylaxis are still controversial in view of sparse data regarding the aviation thromboembolism especially during pregnancy.

Current available guidelines, as published by the British Thoracic Society [10], identify three risk-groups under which pregnancy and 0-2 weeks postpartum is stratified as middle risk; while the American College of Chest Physicians [11] distinguishes 1 increased risk group. In summary, both guidelines recognize high-risk groups and validate their need for thromboprophylaxis. While data comparing mechanical and medical prophylaxis remains scarce, it can be reasoned that Low Molecular Weight Heparins (LMWHs) is to be preferred over below knee stockings for it is a well established prophylaxis. Further more, stockings only work properly in combination with leg muscle movement.

The Green top guideline recommends that women should have their risks discussed and the reasons for individual recommendations explained, citing weight-adjusted dose of LMWHs as the agents of choice for antenatal and postnatal thromboprophylaxis [18]. Their role as the preferred prophylaxis of air-travel-related thrombosis is based on its ability to prevent coagulation activation in a hypobaric chamber [17]. While practices vary in different countries, LMWH by self-injection is the preferred anticoagulant due rapid onset and long-haul coverage (half life ranging 4-6hours post SC administration [19]). A prophylactic dose (0.5mg/kg), one hour before take off, is advisable in pregnant patients with one more risk factors for thrombosis (including height <150cm or >190cm). However, there is not much evidence that recommends the duration of the treatment, although theoretically the risk of VTE persists up to eight weeks post travel.

The risk of bleeding in LMWH usage is noteworthy, as this could be an even bigger hazard when travelling. Studies have shown the risk of major bleeding in LMWH users is 0.14% (95% CI 0.04-0.50%; 2 cases out of 1475 treated [12-15]) on treatment for several weeks. This risk must be substantially lower for a 1-2 days treatment course (estimated 10-fold), assuming the average treatment period is 10 days, while travellers will only be treated for 1 day. Data shows that the risks of VTE outweigh that of bleeding, hence the imperative for thromboprophylaxis in at risk mothers.

Novel oral anticoagulants like Rivaroxaban, Dabigatran, and Apixaban have garnered much popularity. A meta-analysis of several trials shows Dabigatran to be non-inferior to LMWH for VTE prophylaxis [20] with a reduced risk of major hemorrhage. However, in 2014 the ACOG reported a study showing direct ex vivo evidence of Dabigatran and its prodrug crossing a term human placenta [21] rendering this drug best avoided in pregnancy.
4.0 CONCLUSION

Undoubtedly, air travel in at-risk pregnant women significantly increases the risk of VTE yet the proper preventive guidelines are lacking. Mechanical methods alone are insufficient. However, the role of injectable thromboprophylaxis for high-risk women is still debatable due to its method of administration, potential side effects and the efficacy to prevent the thrombo-embolic event. Oral anticoagulants still remain unadvisable to be used in pregnancy.

REFERENCES


[18] Green top guideline no 37a: Reducing the risk of venous thromboembolism during pregnancy and puerperium; April 2015.

