A 39-year-old Rhesus-positive mother had been well. She had been screened low risk (1:2496) for Down syndrome (DS) at the first-trimester combined screening in late 2012. The fetal morphology scan at 20 weeks of gestation was normal. Nevertheless, an ultrasound scan at 32 weeks of gestation showed bilateral pleural effusions. Amnioreduction and left pleural tap yielded 35 mL of pleural chyle. There was no evidence of a viral infection. A girl was delivered at 33-weeks-6-days of gestation by emergency caesarean section because of recurrent fetal pleural effusions. The neonate was mildly oedematous with a distended abdomen and hepatomegaly (Fig 1). The baby was intubated and transferred to the neonatal intensive care unit (ICU) for further management. Pleural effusions (Fig 2) were drained by pleural tapping and chest drains. Echocardiography showed that her cardiac structure and function appeared normal, but a high pulmonary pressure and a patent ductus arteriosus were evident. Plasma total protein was 35 g/L (reference range, 65-82 g/L) and albumin 22 g/L (reference range, 35-52 g/L). The highest blood white cell count was 84.2 x 10^9 /L (50% blasts). What is the underlying diagnosis for this infant’s chylothoraces, hypoproteinaemia, and leukocytosis?

1. Immune hydrops
2. Trisomy syndrome
3. Congenital infection
4. Congenital lymphoma
5. Inborn error of protein metabolism

In the neonatal ICU, she improved with full intensive support (mechanical ventilation, thoracostomy drainage of pleural fluids, and treatment with intravenous octreotide). Trisomy 21 (47,XY,+21) was subsequently confirmed following chromosomal evaluation.

Hydrops fetalis (fetal hydrops) is a serious fetal condition defined as an abnormal accumulation of fluid in two or more fetal compartments, and includes ascites, pleural effusion, pericardial effusion, and skin oedema.\(^1\) It may be due to immune or non-immune aetiologies.\(^1,2\) Rhesus isoimmunisation is the commonest immune aetiology, and alpha-thalassaemia is a non-immune cause.\(^1\) Immediate diagnosis of other aetiologies is often not possible without extensive investigations. A prompt spot diagnosis of DS was made in this neonate with typical facial features, which obviated the need for an extensive search for an underlying aetiology and enabled target therapies to be instituted (Fig 1). Trisomy 21 is a known association with hydrops fetalis and myeloproliferative disorder.\(^1,2\)

Transient myeloproliferative disorder (TMD) is a self-limiting disorder characterised by leukocytosis and the presence of megakaryoblasts in the peripheral blood and bone marrow, anaemia, thrombocytopenia, and organomegaly. It occurs in...
approximately 10% of newborn infants with DS.\(^2\)

Hepatic fibrosis is encountered in the severe form of TMD with DS, and is characterised by diffuse intralobular sinusoidal fibrosis and extramedullary haematopoiesis.\(^3\) Although TMD in most patients resolves spontaneously within the first 3 months of life, in a few severe cases there can be hepatic fibrosis or cardiopulmonary failure. Acute megakaryocytic leukaemia (AML-M7) is noted in 20 to 30% of babies with DS and TMD within the first 4 years of life.\(^4\) Cytokines produced by megakaryocytes (including transforming growth factor-beta, platelet-derived growth factor, and platelet factor 4) could be responsible for the pathogenesis of TMD.\(^3\) The imbalance between intravascular or capillary hydrostatic pressure and transcapillary filtration may be responsible from hydrops fetalis.\(^2,5\)

Prompt recognition of the facial features of DS is important to facilitate immediate diagnosis and management of this neonate with hydrops fetalis.

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