

Background

Immune anti-D

This category was introduced in 2012 as a separate study from the standard SHOT reporting categories. Accordingly, SHOT has been reviewing cases where immune anti-D has been detected for the first time, in the current pregnancy, to improve understanding of the causes of continuing anti-D immunisations. The cases reported under this category are not included in the total number of SHOT reports analysed in the Annual SHOT Report.

Anti-D Ig errors

Events relating to administration of anti-D immunoglobulin (Ig) were included since the 1997/98 Annual SHOT Report. At present, this category includes adverse events relating to the requesting and/or administration of anti-D Ig during pregnancy or after delivery, and adverse events relating to administration of anti-D Ig to D-negative patients with childbearing potential including paediatric, following transfusion of D-positive blood components and following D-mismatch solid organ transplants.

Basic principles of Anti-D Ig errors and Immune anti-D

- Two different SHOT categories
- Two different SHOT definitions (see resources)
- Two different purposes
- Two different remits
- Managed differently by SHOT
- Might need two separate SHOT reports



Reactions to anti-D Ig are **not reportable to SHOT** but they are reportable to the Medicines and Healthcare Regulatory Agency (MHRA) via the 'Yellow Card' scheme for medicines (<https://yellowcard.mhra.gov.uk/>)



There are cases where the adverse event needs to be reported to both categories (two SHOT reports). These include cases where:

*Event relating to requesting and/or administration of anti-D Ig during pregnancy or after delivery e.g., late or omitted anti-D Ig administration, wrong dose, false negative cffDNA screening result → **Anti-D***

AND

*Alloimmunisation detected during pregnancy → **Immune anti-D***

Examples of the different remit for both categories

Is this event SHOT reportable?	Immune anti-D	Anti-D
Patient non-compliance resulting in immune anti-D	Yes	No
No error but patient found to have immune anti-D	Yes	No
Error with anti-D Ig administration but no D sensitisation	No	Yes
Error with anti-D Ig administration and D sensitisation confirmed	Yes	Yes
Sensitisation identified in pregnancy but reason unknown	Yes	Variable

Errors relating to requesting and/or administration of anti-D Ig during pregnancy or after delivery can cause D sensitisation

SHOT analysed **2963** incidents with potential to cause D sensitisation in D-negative women of childbearing potential due to omission or late administration of anti-D Ig (2012-2022)

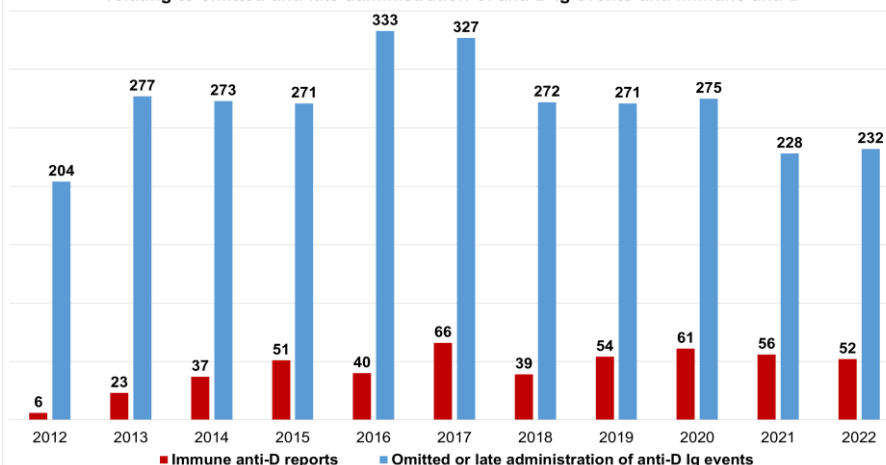
Incorrect management of D-negative pregnancies with D-positive fetus including omission or late administration of anti-D Ig can result in **MAJOR MORBIDITY** 'Sensitisation to D or K in a woman of childbearing potential' (SHOT definitions, 2024)

SHOT reports relating to immune anti-D analysed from 2012 to 2022 (n=485)

Common themes

- Ideal management does not equal no sensitisation
- Delivery beyond 40 weeks may be a risk factor for sensitisation even when managed appropriately
- A postpartum fetomaternal haemorrhage >4mL may be a risk factor for sensitisation even when managed appropriately
- Women who are obese may not be adequately protected by standard doses of anti-D Ig
- There are missed opportunities where pregnancy management is not ideal

Comparison of the number of reports analysed by SHOT between 2012 and 2022 relating to omitted and late administration of anti-D Ig events and immune anti-D



Other contributory factors:

- Mother D-status: D-variant
- cffDNA screening testing (false negative)
- Lack of IT systems interoperability (clinical and laboratory IT systems)
- Anti-D Ig errors
- Gap in knowledge
- Misinterpretation of results



Remember: Report all cases of alloimmune anti-D found for the first time in pregnancy to SHOT. Please provide a complete data set relating to index and previous pregnancies to help identify and understand risk factors for D sensitisation.

The impact of immune anti-D in pregnancies: the numbers (2018-2022)

Outcome of index pregnancy

4 miscarriages

230 live births

2 stillbirths

2 terminations

19 cases
No information*

4 intrauterine deaths

Number of cases where treatment was required for signs and symptoms of haemolytic disease of the fetus and newborn

- ✓ Phototherapy (n=59)
- ✓ Exchange transfusion (n=8)
- ✓ Phototherapy and intravenous immunoglobulin (IVIg) (n=4)
- ✓ Phototherapy and exchange transfusion (n=6)
- ✓ Phototherapy and intrauterine transfusion (IUT) (n=1)
- ✓ IUT and transfusion after delivery (n=1)
- ✓ Phototherapy, IVIg and exchange transfusion (n=4)

*It is important that all details from index (current) and previous pregnancies are reported to help identify and understand the risks and causes for D sensitisation

Useful and relevant resources

- ✓ Anti-D Ig Administration in Pregnancy- an aide memoire: <https://www.shotuk.org/resources/current-resources/>
- ✓ BSH guideline for the use of anti-D immunoglobulin for the prevention of HDFN: <https://doi.org/10.1111/tme.12091>
- ✓ BSH guideline for the Estimation of FMH: <https://b-s-h.org.uk/guidelines/guidelines/the-estimation-of-fetomaternal-haemorrhage>
- ✓ IBGRL Fetal screening: <https://www.nhsbt.nhs.uk/ibgrl/services/molecular-diagnostics/fetal-rhd-screen/>
- ✓ IBGRL Fetal genotyping – diagnostic: <https://www.nhsbt.nhs.uk/ibgrl/services/molecular-diagnostics/fetal-genotyping-diagnostic/>
- ✓ NICE guideline [NG201] – Antenatal care: <https://www.nice.org.uk/guidance/ng201/chapter/Recommendations>
- ✓ IT supports anti-D Ig management in pregnancy: <https://www.shotuk.org/resources/current-resources/>
- ✓ SHOT Bite No 2: Anti-D Ig Administration: <https://www.shotuk.org/resources/current-resources/shot-bites/>
- ✓ SHOT video Anti-D Ig and Immune anti-D (part 1 and part 2): <https://www.shotuk.org/resources/current-resources/videos/>
- ✓ SHOT definitions 2024: <https://www.shotuk.org/reporting/>

