

## Naranjo Nomogram for Adverse Drug Reaction Assessment

### Probability

Definite: Score  $\geq 9$

Probable: 5-8

Possible: 1-4

Doubtful:  $\leq 0$

(Michel DJ, Knobel Lc. Comparison of three algorithms used to evaluate adverse drug reactions. Am J Hosp Pharm. 1986; 43:1711)

Case 1 by: <b>Torrente et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Rapid improvement after corticosteroids and azathioprine administration Biopsy conformation of AIH
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	0	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 5→ Probable adverse drug reaction					

Case 2, 3, 4, 5 by: <b>Izagirre et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	$\geq 2$ published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Rapid improvement after immunomodulators administration for 3 of the patients. One of the patients did not need any drug remedy. 3 patients had booster dose without any complaints  Biopsy conformation of AIH for 3 of the patients. One patient did not have liver biopsy.
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	Score may be 0 to 1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	Score may be (-)1 to 0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	0	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	Score may be 0 to 1	
Total score may be from <b>2 to 5</b> for each patient $\rightarrow$ <b>Possible/Probable adverse drug reaction</b>					

Case 6 by: <b>Barary et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	<div>≥2 published reports in which the adverse reaction has been described in detail</div> <div>Adverse reaction appeared soon after vaccination</div> <div>No improvement after high-dose corticosteroids administration.</div> <div>The patient was treated with hepatotoxic agents for psychiatric disease</div> <div>Biopsy was not compatible with AIH</div>
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	0	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	-1	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	0	
Total score: 2 →Possible adverse drug reaction					

Case 7 by: <b>Mekritthikrai et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Rapid improvement after immunomodulator therapy
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could provoke acute hepatitis
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	Biopsy typical for AIH
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 8 by: <b>Brubaker et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	0	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could flare of AIH
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	Elevation of ASMA and hepatic enzymes. Biopsy confirmed AIH in the past.
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 6 → Probable adverse drug reaction					

Case 9 by: <b>Shahrani et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could lead to AIH
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	Biopsy compatible with AIH
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 10 by: <b>Shahrani et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Temporary improvement with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could lead to AIH
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	Biopsy typical for AIH
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 11 by: <i>Shahrani et al</i>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could lead to AIH
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 12 by: <b>Hasegawa et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Rapid improvement with immunomodulators
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could lead to AIH
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy typical for AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 13 by: <i>Lasagna et al</i>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Rapid improvement with immunomodulators
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	-1	Cancer patient on immunotherapy
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy non-consistent with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	0	
Total score: 3 → Possible adverse drug reaction					

Case 14 by: <b>Pinazo-Bandera et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with immunomodulators
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could lead to AIH
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy consistent with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 15 by: <b>Pinazo-Bandera et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with immunomodulators  No known agent or condition that could lead to AIH
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 16 by: <b>Boettler et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with corticosteroids  No known agent or condition that could lead to AIH
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	2	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 9 → Definite adverse drug reaction					

Case 17 by: <b>Kang et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with corticosteroids  No known agent or condition that could lead to AIH
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 18 by: <i>Ghorbani et al</i>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	0	No known agent or condition that could lead to AIH
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 6 → Probable adverse drug reaction					

Case 19 by: <b>Fimiano et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with corticosteroids No known agent or condition that could lead to AIH
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 20 by: <i>Camacho-Dominguez et al</i>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with corticosteroids No known agent or condition that could lead to AIH
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Cases 21,22 by: <b>Erard et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with corticosteroids No known agent or condition that could lead to AIH
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 23 by: <i>Erard et al</i>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination No improvement with immunomodulators No known agent or condition that could lead to AIH
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	0	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 6 → Probable adverse drug reaction					

Cases 24, 25, 26 by: <b>Suzuki et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 27 by: <b>Cao et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 28 by: <b>Avci et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 29 by: <b>Garrido et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 30 by: <b>Palla et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 31 by: <b>Zhou et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	Patient had a diagnosis of primary sclerosing cholangitis (PSC). However, she was on remission under continuous monitoring and flare of PSC is considered unlikely.
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 32 by: <b>Ghielmetti et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 33 by: <b>McShane et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 34 by: <b>Clayton-Chubb et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	Biopsy compatible with AIH
Total score: 7 → Probable adverse drug reaction					

Case 35 by: <b>Tan et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	Biopsy compatible with AIH
Total score: 7 → Probable adverse drug reaction					

Case 36 by: <b>Rocco et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	Biopsy compatible with AIH
Total score: 7 → Probable adverse drug reaction					

Case 37 by: <b>Bril et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Improvement with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	Biopsy compatible with AIH
Total score: 7 → Probable adverse drug reaction					



Case 38 by: <b>Goulas et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	Adverse reaction appeared soon after vaccination
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	Improvement with corticosteroids
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could lead to AIH
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	Biopsy compatible with AIH
Total score: 7 → Probable adverse drug reaction					

Case 39 by: <b>Lodato et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	Adverse reaction appeared soon after vaccination
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	Resolution with corticosteroids
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could lead to AIH
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	Biopsy compatible with AIH
Total score: 7 → Probable adverse drug reaction					

Case 40 by: <b>Londono et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	Adverse reaction appeared soon after vaccination
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	2	Resolution with corticosteroids
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	Precipitation of symptoms after second vaccination
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	No known agent or condition that could lead to AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	Biopsy compatible with AIH
Total score: 9 → Definite adverse drug reaction					

Case 41 by: <b>Rela et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	Adverse reaction appeared soon after vaccination
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	Resolution with corticosteroids
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	No known agent or condition that could lead to AIH
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	Biopsy compatible with AIH
Total score: 7 → Probable adverse drug reaction					

Case 42 by: <i>Rela et al</i>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Poor response to corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	0	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	Medical history of jaundice episodes
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	-1	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 3 → Possible adverse drug reaction					

Case 43 by: <b>Zin Tun et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	2	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: <b>9</b> → Probable adverse drug reaction					

Case 44 by: Vuille-Lessard et al	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with immunomodulators
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 7 → Probable adverse drug reaction					

Case 45 by: <b>Mathew et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with immunomodulators
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	Chronic use of analgetics. Possible drug toxicity is described also in liver histopathological findings.
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	-1	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 4 → Possible adverse drug reaction					

Case 46 by: <b>Ferronato et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Spontaneous resolution
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	0	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appear when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 6 → Probable adverse drug reaction					

Case 47 by: <b>Ferronato et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Patient declined corticosteroids. Symptoms persist.
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	0	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	No known agent or condition that could lead to AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appear when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 6 → Probable adverse drug reaction					

Case 48 by: <b>Ferronato et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Flare of transaminases elevation appeared soon after vaccination Resolution with corticosteroids.
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	Known AIH, but symptoms flare with vaccination
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appear when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Flare was more severe after 2 <sup>nd</sup> vaccine dose
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	1	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	Biopsy compatible with AIH
Total score: 8 → Probable adverse drug reaction					

Case 49 by: <b>Romero-Salazar et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with corticosteroids.
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	Patient with PBC, but liver biopsy was typical for AIH
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appear when a placebo was given?	-1	+1	0	0	
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	Biopsy compatible with AIH
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	

Case 50 by: <b>Mahalingham et al</b>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution with immunosuppression.
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	1	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	Patient with liver transplantation due to AIH, but no evidence of active disease before vaccination.
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: <b>7</b> → Probable adverse drug reaction					

Case 51 by: <i>Lee et al</i>	Yes	No	Do not know	Score	Comment
1. Are there previous conclusive reports on this reaction?	+1	0	0	1	≥2 published reports in which the adverse reaction has been described in detail Adverse reaction appeared soon after vaccination Resolution without need for immunomodulation.
2. Did the adverse event appear after the suspected drug was administered?	+2	-1	0	2	
3. Did the adverse reaction improve when the drug was discontinued, or a specific antagonist was administered?	+1	0	0	0	
4. Did the adverse reaction reappear when the drug was readministered?	+2	-1	0	0	
5. Are there alternative causes (other than the drug) that could on their own have caused the reaction?	-1	+2	0	2	
6. Did the reaction appeared when a placebo was given?	-1	+1	0	0	No known agent or condition that could lead to AIH
7. Was the drug detected in the blood (or other body fluids) in concentrations known to be toxic?	+1	0	0	0	
8. Was the reaction more severe when the dose was increased, or less severe when dose decreased?	+1	0	0	0	
9. Did the patient have a similar reaction to the same or similar drugs in any previous exposure?	+1	0	0	0	
10. Was the adverse event confirmed by any objective evidence?	+1	0	0	1	
Total score: 6 → Probable adverse drug reaction					