Australian Government
Department of Health
Therapeutic Goods Administration

Electronic submission of individual case safety reports

Electronic data interchange for ICSR submission using the E2B R2 format

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TGA Health Safety Regulation

Contents

Providing information about adverse events using E2B (R2) messages 5

Data quality principles of ICSRs 7

Minimum information 7

Format of data elements 8

Registration for E2B R2 adverse event reporting 9

Connecting to the EDI service 9

Generating a valid ICH safety message 10

ICSR business rules 11

M.1 ICH ICSR message header specifications 11

A. E2B (R2) ICSR specifications 11

B. Information on the case 11

Use of dates and intervals 11

B.1.7 Relevant medical history and concurrent conditions 12

B.1.8 Relevant past drug history 12

B.2 Reaction(s) or event(s) 12

B.3 Results of tests and procedures relevant to the investigation of the patient 12

B.4 Drug(s) information 12

Acknowledgment of reports 15

ICSR acknowledgement message 17

Updating previously submitted ISCRs 18

Duplicate reports 18

Privacy and data security 19

Personal information of sender and primary source 19

Privacy statement 19

Data security 20

TGA specific validation rules 21

M.1 ICH ICSR message header specifications 21

ICH ICSR M2 data processing specifications ---------- 22
A. E2B (R2) ICSR specifications _____ 23
A.1 Safety report ---------- 23
A.2 Primary source(s) of information ---------- 28
A.3 Information on sender and receiver of ISCR ---------- 31
B. Information on the case _____ 35
B.1 Patient characteristics ---------- 35
B.2 Reaction(s) or event(s) ---------- 50
B.3 Results of tests and procedures relevant to the investigation of the patient ---------- 53
B.4 Drug(s) information ---------- 54
B.5 Narrative case summary and further information ---------- 61
Test scenarios to use the E2B standard _____ 62
Tests _____ 62

Providing information about adverse events using E2B (R2) messages

The E2B (R2) EDI service is intended for use by regulated industry and regulators to report ICSRs relating to medicines and vaccines only. Adverse events that occur during clinical trials may also be reported using the E2B (R2) EDI service.

The E2B (R2) standard does not accommodate medical device adverse event (incident) reports. Therefore, sponsors of medical devices can submit their reports via the Sponsors/Manufacturers medical device incident report page.

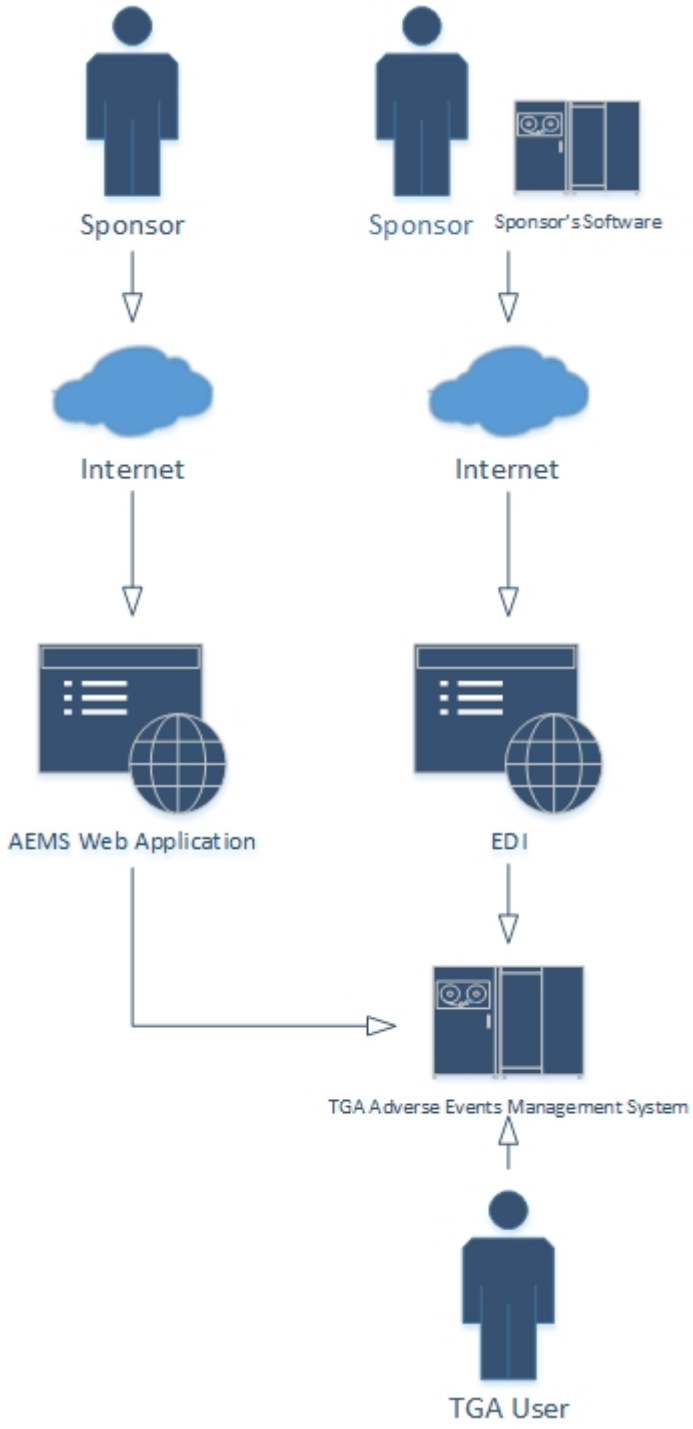
The TGA can receive information on adverse events from regulated industry and regulators using E2B (R2) messages in the Extensible Markup Language (XML) format by Electronic Data Interchange (EDI). This implementation guide specifies how your systems can submit individual case safety reports (ICSRs) for medicines including vaccines to the TGA using the E2B (R2) EDI service.

The preparation of an E2B (R2) file requires mapping of your internal database to the E2B (R2) data set described in the ICH M2 EWG Electronic Transmission of Individual Case Safety Reports Message Specification (ICH ICSR DTD). The ICH ICSR DTD defines the specific data elements to be included for electronic submission of adverse event data, the order of the elements, and their interrelationships.

After your database has been mapped, the transmitted data must be in conformance with the ICH ICSR DTD and the TGA specific validation rules outlined in this guidance.

The TGA also has a separate web based reporting system that you can use to continue reporting adverse events while you set up your systems to submit E2B (R2) reports using the E2B (R2) EDI service or if your systems are not compatible with this format. The following diagram illustrates the two pathways for submitting ICSRs.

Figure 1: Pathways for submitting ICSRs



AEMS: Adverse Events Management System

Data quality principles of ICSRs

Sponsors must ensure that all information relevant to the balance of benefits and risks of a registered or listed medicine is reported to the TGA in accordance with Australian requirements and recommendations for pharmacovigilance responsibilities of sponsors of medicines, including medical and administrative data related to the adverse event.

ICSRs should include the complete information for an individual case that is available to the sender. This information must be entered in a fully structured format using all applicable and relevant ICH E2B (R2) data elements and terminologies, which should be repeated as necessary. This applies to all types of ICSRs, that is:

- initial information on the case
- reports with follow-up information
- cases highlighted for nullification – ensure that:
  - A.1.13: 'Report nullification' is set to 'yes'
  - A.1.13.1: 'Reason for nullification' is completed.

Associated documents cannot be submitted using the TGA E2B functionality. Any supporting information related to the individual case should be sufficiently described within A.1.8.2: 'List of documents held by sender' and only provided upon request by the TGA. For adverse event reports from literature, the literature reference must be provided in A.2.2 'Literature reference(s)'and the abstract included in B.5.1 'Case Narrative' if possible.

Any information concerning previous transmissions of the same case from either other senders or by the same sender using a different reporting channel such as email or online form should be provided in the A.1.11 'Other case identifiers in previous transmissions'. This is to aid the detection and management of duplicates.

Date information within the ICSR should follow a logical sequence. For example, for the data element 'Date of start of reaction/event' (B.2.i.4b) it would be expected that the supplied date:

- does not refer to a future date
- is less than or equal to the date of transmission (A.1.3b)
- is greater than or equal to the date of birth of the patient (B.1.2.1b)
- is greater than or equal to the start date (B.4.k.12b) for at least one drug with a drug characterisation role of 1 (suspect) or 3 (interacting).

Minimum information

While complete information is desirable, a minimum set of information is always required for an ICSR to be valid. This applies to all types of ICSRs including initial case reports, and cases to be amended or nullified. Minimum data requirements for ICSR sent to the TGA, including mandatory and conditionally mandatory fields are specified in the TGA specific business rules.

Format of data elements

All the information available should be reported in a fully structured format using the relevant E2B (R2) data elements, applicable standard terminologies and TGA specific business rules.

Standard terminologies used in ICSRs include:

- ISO (country codes (2 character), gender codes and language codes)
- Medical Dictionary for Regulatory Activities (MedDRA)
  - Where MedDRA terms are used (for example, medical history, indication, and reaction / event), the same MedDRA version must be used throughout the ICSR.
- UCUM (units of measurement)
- ICH M2 (for example, units and routes of administration).

Please refer to each standard for further information.

Registration for E2B R2 adverse event reporting

You will need to carry out testing with the TGA to ensure that you meet the requirements and that your systems are compatible.

You need to follow this process to start using the E2B (R2) EDI service:

1. Familiarise yourself and ensure your system complies with the following formatting standards:
   - ICH ICSR DTD
   - TGA specific validation rules specified.
   - Coding standards specified by the Medical Dictionary for Regulatory Activities.
2. Complete the registration process. The registration process will include:
   - Emailing us to register your interest in E2B R2 adverse event reporting.
   - The exchange of technical and contact details using TGAs template form.
   - The exchange of relevant digital certificates used for authentication and signing.
   - The nomination of earliest and preferred dates for testing.
   - The establishment of a new connection in the TGA system for your organisation.
   - The establishment of a new connection for TGA in your system.
3. Successfully transmit test cases to the test E2B (R2) EDI service after establishment of the connection:
   - The required test scenarios for sponsors to complete are described below.
   - Send an email after you have submitted the test scenarios.
     - We will advise you on the outcome of the testing.
   - Contact TGA via email if you encounter any issues during testing.
4. Following successful testing, you will be able to use the production E2B (R2) EDI service.

You can continue to send adverse event reports using the TGA online reporting form while you set up for E2B (R2) reporting.

Connecting to the EDI service

Following registration and testing of your systems, you can begin submitting ICSRs to the TGA production E2B (R2) EDI service. The following sections describe the specifications for generating and submitting ICSRs.

Each ICH Safety Message should only contain a single adverse event report. Batches of adverse event reports in the same ICH Safety Message should not be submitted. There is a size restriction on the xml file of 2 megabytes.

Generating a valid ICH safety message

There are two levels of conformance in the XML specifications: a well-formed and a valid message.

1. A well-formed message is an XML document that conforms to the structural rules of XML:
   - The first line should be the XML document declaration
   - The document should contain at least one element (or tag)
   - Every starting tag should have a closing tag
   - Self-closing tags (for example, '<tag/>') are permitted when XML elements do not contain data
   - Tags cannot overlap.

   In addition, as XML is case sensitive, all the fields and attribute names have to be in lower case in order to comply with the XML DTD.

2. A Valid XML file is one, which has a DTD reference and which conforms to the ICH ICSR DTD and the TGA ICSR business rules.

   The ICH ICSR DTD defines the valid elements (tags), attributes that may appear in a particular type of XML document and also defines element nesting rules.

   The following XML special characters (excluding quotation marks) ">", "<" and "&" when occurring in text should always be replaced by ">" "<" and "&" respectively. Regarding all aspects of XML, the W3C standards should be followed.

ICSR business rules

Follow the ICH ICSR DTD and TGA specific validation rules when sending ICSRs to TGA and to correct any verification error messages.

The alphanumeric character limits, codes and formatting rules for each field must be adhered to, as specified in the A.1 ICSR Attribute List of the ICH ICSR DTD to ensure transmission of the reports.

Where dates are provided, the date should be on or before the transmission date. Future dates must not be used.

M.1 ICH ICSR message header specifications

A Safety Message sent to the TGA must contain only one ICSR. The ICSR contains a Message Header part which should include information on the sender, the receiver, the message date and a unique message identification number.

A. E2B (R2) ICSR specifications

Section A provides information on the following aspects of the ICSR:

- A.1 Safety report
- A.2 Primary source(s) of information
- A.3 Information on sender and receiver of ISCR.

B. Information on the case

Section B provides the following information on the adverse event:

- B.1 Patient characteristics
- B.2 Reaction(s) or event(s)
- B.3 Results of tests and procedures relevant to the investigation of the patient
- B.4 Drug(s) information
- B.5 Narrative case summary and further information.

Use of dates and intervals

When specifying the duration relating to an adverse event, the preferred format is for dates to be transmitted in the appropriate items, rather than intervals.

Intervals may be used where only imprecise dates are known but more information concerning the interval is known. For example, where there is a very short interval between drug administration and start of reaction or event, such as in anaphylaxis.

Dates specified in ICSRs cannot refer to a future date. Any future dates will generate an error message during validation.

B.1.7 Relevant medical history and concurrent conditions

Medical judgment should be exercised in completing this section. Information pertinent to understanding the case is desired, such as diseases, conditions such as pregnancy, surgical procedures, and psychological trauma. Each of the items in the section can be repeated as appropriate.

Information can be included as comments if precise dates are unknown and a text description would aid in understanding the medical history, or if concise additional information is helpful in showing the relevance of the past medical history.

B.1.8 Relevant past drug history

This section concerns drugs previously taken, but not those taken concomitantly or drugs that may have potentially been involved in the current reaction or event. Information concerning concomitant and other suspect drugs should be included in section B.4.

The information provided in B.1.8 can also include previous experience with similar drugs. Medical judgment should be exercised in completing this section.

B.2 Reaction(s) or event(s)

The designation of *i* in this section indicates that each item is repeatable and that it carries an appropriate correspondence to the same *i* in all subsections. A separate block (*i*) should be used for each reaction or event term. For example, if two reactions are observed, the first reaction would be described in items B.2.1.0 through B.2.1.8, and the other reaction would be described in items B.2.2.0 through B.2.2.8. The reaction or event specified in the first iteration should be the one used in assessing the intervals in B.4.k.13.

B.3 Results of tests and procedures relevant to the investigation of the patient

This section should capture the tests and procedures performed to diagnose or confirm the reaction or event, including those tests done to investigate (exclude) a nondrug cause, (for example, serologic tests for infectious hepatitis in suspected drug-induced hepatitis). Both positive and negative results should be reported where known. Although structured information is preferable, provisions have been made to transmit the information as free text in B.3.2.

B.4 Drug(s) information

This section covers both suspect drugs (including all interacting drugs) and concomitant medications, including vaccines. For each drug, the characterisation of the drug role (B.4.k.1) is that indicated by the primary reporter (the original source of the information).

The designation *k* in this section indicates that each item is repeatable and that it carries an appropriate correspondence to the same *k* in all subsections. A separate block (*k*) should be used for each drug. The drug specified in the first iteration should be the one used in assessing the intervals in data element B.2.i.7. Drugs used to treat the reaction or event should not be included in section B.4.

The Australian approved name should be provided where possible but if unavailable the substance name/s should be provided. In the case of investigational drugs, only a code may be known and provided. For clinical trials involving a product included in the ARTG, the Australian Approved Name should be provided.

If a single drug contains multiple active substances, each should be specified as a repeating data element B.4.k.2.2.

B.4.k.5 Structured dosage information

Table 1 provides examples for completing data elements B.4.k.5.1 through B.4.k.5.7.

The cumulative dose provided is the total dose administered until the first sign, symptom, or reaction. For prolonged chronic therapy, the sender should consider the need to complete the cumulative dose data elements.

In the case of a parent-child or parent-foetus report, the dosage data elements apply to the parental dose.

For dosage regimens that involve more than one dosage form and/or changes in dosage, the information can be provided in data element B.4.k.6 as text. Alternatively, the sender can provide more than one iteration (*k*) for the same drug.

Table 1: Examples of structured dosage information at B.4.k.5

Data element	Example 1 2 mg three times a day for five days	Example 2 5 mg (in one dose) every other day for 30 days	Example 3 50 mg daily for 2 days
B.4.k.5.1 dose (number)	2	5	50
B.4.k.5.2 dose (unit)	mg	mg	mg
B.4.k.5.3 number of separate dosages	3	1	1
B.4.k.5.4 number of units in the interval	1	2	1
B.4.k.5.5 definition of the interval unit	day	day	day
B.4.k.5.6 cumulative dose to first reaction (number)	30	75	100
B.4.k.5.7 cumulative dose to first reaction (unit)	mg	mg	mg

B.4.k.18 Relatedness of drug to reactions or events

This repeatable block provides the means to transmit the degree of suspected relatedness of each drug to the reaction or event. The repeating elements could also be used to provide the assessment of relatedness by different sources or methods of assessment. For the purpose of reporting, there is an implied suspicion of causality for spontaneous reports. It is recognised that information concerning the relatedness, especially for spontaneous reports, is often subjective and may not be available.

Example of extensive functionality

The following example illustrates the extensive functionality contained in this section:

- A patient was being treated with two medications: Drug A and Drug B.
- The patient had two adverse events: Event 1 and, Event 2
- The reporter provided assessment of causality for event 1 for both Drug A and Drug B, but not for event 2 for either drug. The reporter's assessment of causality is based on overall impression, which the sender codes as "global introspection".
- The sender applied causality assessment with an algorithm (coded algorithm), but it does so only for the drug it manufactures (in this case Drug A) for both reported events.
- Therefore, there are 2 sets of data for the reporter (2 drug x1 events x1 method of assessment) and 2 sets for the sender (1 drug x2 events x1 method of assessment) for a total 4 sets of data.
- The appropriate data element for the information is B.4.k.18 (and its four subfields 1-4) as shown in table 2. In this example k is replaced by Drug A and Drug B respectively. Please note the subfields 1-4 are repeatable.

Table 2: Example of extensive functionality at B.4.k.18

B.4.k.18.1	B.4.k.18.2	B.4.k.18.3	B.4.k.18.4
k(1) = Drug A			
Event1	Reporter	global introspection	related
Event1	Company	algorithm	possibly related
Event2	Company	algorithm	possibly related
k(2) = DRUG B			
Event1	Reporter	global introspection	not related

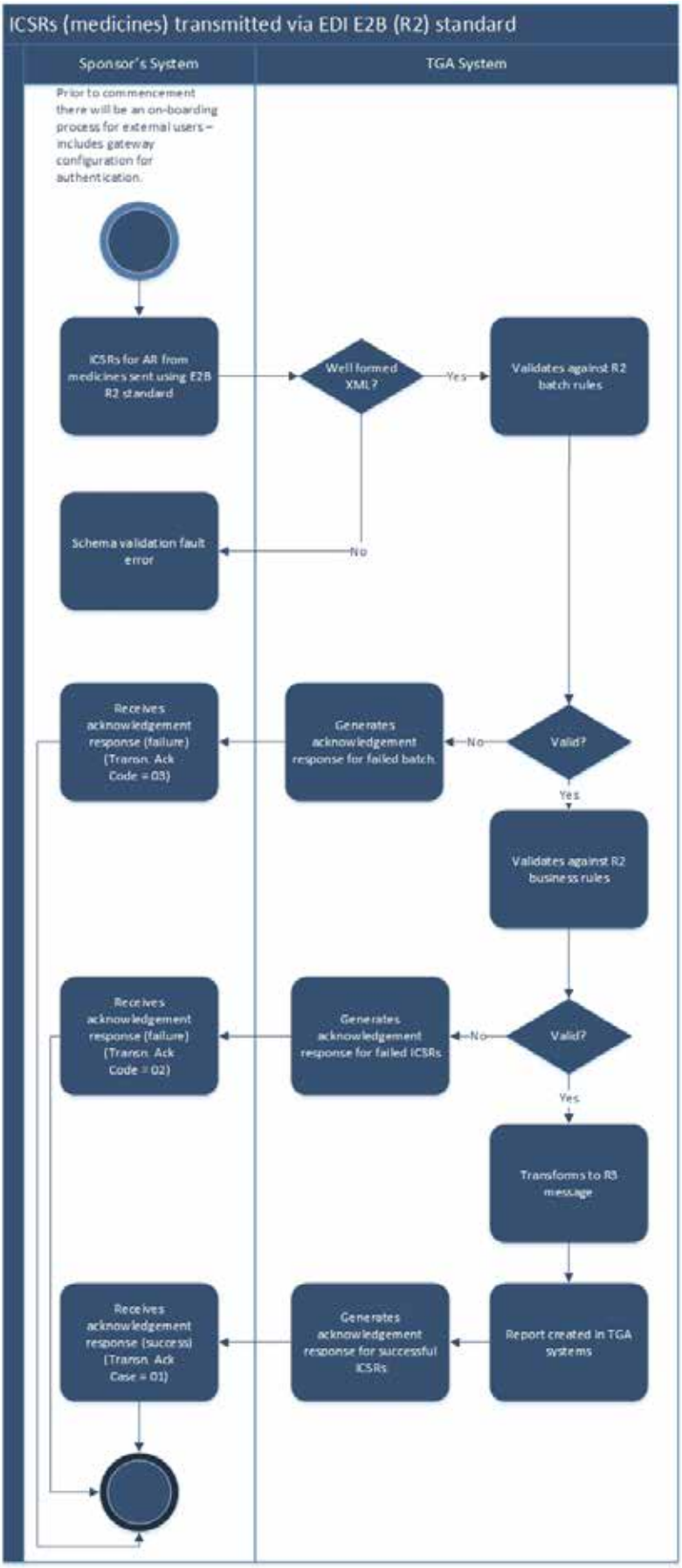
The order of the rows is not important since each one represents a complete set, however the E2B message and M2 specifications state that all assessments for Drug A (k=1) appear before Drug B (k=2).

Acknowledgment of reports

Following submission, you will receive a message delivery notification (MDN) from our system confirming receipt of the message. An acknowledgment message relating to the success or failure of the transmission at both the ICSR batch and ICSR message level will be sent following processing of the message. Validation of ICSRs to the ICSR business rules occurs in the following stages, which are illustrated in figure 1:

1. Batch transmission - formatting compliance of a batch of XML files
3. Message header validation - compliance with the R2 batch rules
4. ICSR validations – compliance of individual cases with the business rules.
5. Acknowledgement of successful transmission.

Figure 2: ICSRs (medicines) transmitted via EDI using the E2B (E2) standard



At each stage, acknowledgment messages may be generated relating the transmission of the report. If a batch message does not have a well-formed XML structure, a general error message, “SchemaValidationFault” will be returned. The XML should be reviewed to ensure it follows the formatting specified in the ICH ICSR DTD.

ICSR acknowledgement message

Following transmission of an ICSR report, you will receive an acknowledgement message. This message advises on whether the batch of ICSRs was successfully transmitted.

The codes are:

01 all ICSR reports in the batch have successfully loaded

02 not all ICSR reports in the batch have successfully loaded – refer to the ICSR business rules.

03 batch validation error, no ICSR reports in the batch have successfully loaded – refer to the ICSR business rules.

This message also advises whether the individual ICSR was successfully processed. The codes are:

01 report loaded successfully

02 report not loaded – refer to the ICSR business rules.

When a report is not loaded the acknowledgment message will include information on the first encountered error in B.1.9 errormessagecomment. The default error message is as follows:

<<DTD Descriptor>>:'Invalid data supplied. Please refer to the TGA EDI ICSR R2 Implementation Guidelines'.

Certain errors will result in specific error messages which are detailed below in the TGA specific validation rules for the relevant data elements.

All cases that are not loaded should then be corrected in the identified field to align with the ICH ICSR DTD and the ICSR business rules, which includes guidance for correcting issues identified in acknowledgement messages. The corrected case/s should then be re-transmitted to the E2B R2 EDI service.

Note: Batches sent to the TGA must contain only one ICSR. Where the ICSR did not load, the acknowledgment message will only provide information on the first error encountered in sequential order of the DTD element number as defined in the R2 standard. Therefore, additional errors may be identified in subsequent validations of the same case.

Updating previously submitted ISCRs

ISCRs may be updated following the initial transmission of an ICSR by submitting a new version of the ICSR. The process for submitting an updated version of the ICSR follows the same process as for the original case, however there are some additional validation rules that apply when submitting an updated version of an ICSR. These are specified in the ICSR business rules. Amended ISCRs submitted with a repeated version number and transmission date will be detected as a duplicate and will not be loaded.

The TGA uses a classification process to manage the versioning of the incoming ICSRs and maintain the entire history of the case reports related to a specific ICSR.

A report may be classified as:

- Initial report
  - a report describing a case for the first time
- Amended report
  - a report describing a case at a later time
- Nullified report
  - Nullified Report is a report with the data element *casenullification* (ICH E2B (R2) A.1.13) set to 'yes'.

Note: After a case has been nullified it can no longer be updated. To resubmit, the ICSR should be submitted as a new ICSR.

Duplicate reports

To assist with managing duplicates the E2B R2 web service undertakes checks using the Sender's (Case) Safety Report Unique Identifier (A.1.0.1) and the Worldwide unique case identification number (A.1.10) to identify duplicate reports. These checks are specified in the ICSR business rules.

If you are amending a case which was previously sent to the TGA via a different reporting channel such as email or online form, you will need to include the other case identifier details in A.1.11.

Privacy and data security

Personal information of sender and primary source

The TGA does not require personal information on the individual in the Sender and Primary Source entities to be included in the ICSR.

Specifically, the following fields are not required to have a value:

- sendertitle (A.3.1.3b)
- sendergivename (A.3.1.3c)
- sendermiddlename (A.3.1.3d)
- senderfamilyname (A.3.1.3e)
- reportertitle (A.2.1.1a)
- reportergivename (A.2.1.1.b)
- reportermiddlename (A.2.1.1c)
- reporterfamilyname (A.2.1.1d)

The E2B R2 EDI service will not store any values from the above fields (if they are provided) within the TGA system.

Privacy statement

For general privacy information, go to the TGA Privacy web page.

Individual Case Safety Reports (ICSRs) are collected to assist in the post market monitoring of the safety of therapeutic goods under the *Therapeutic Goods Act 1989*. All reports are entered into the Therapeutic Goods Administration's (TGA) Australian Adverse Event Management System. Further information about how the TGA uses adverse event information that is reported to it is available at the TGA Reporting Adverse Events web page.

The TGA collects personal information in ICSRs to:

- monitor the safety of medicines and vaccines under the Act
- contact the sender of the adverse event if further information is required
- contact representatives of entities that supply therapeutic goods, to discuss reported adverse events
- check that the same information has not been received multiple times for the same adverse event.

At times, this information is collected from someone other than the individual to whom the personal information relates. This can occur when an adverse event is reported to a person or an entity other than the TGA (such as a health professional or a hospital), and that person or entity passes the information on to the TGA.

Personal information collected in this report may be disclosed by consent or where the disclosure is required by, or authorised under, a law (for example, under section 61 of the Act). Where a report relates to vaccine events, personal information about the reporter or the patient may be disclosed to State and Territory health agencies under subsection 61(3) of the Act.

It is the TGA's preference that personal information, such as the names of patients, health professionals, or health facilities; date of birth; and patient identification numbers, are not included in the case narrative. This information should instead be provided in the sender comments or other applicable fields.

Data security

Messages sent to and received from the service will be encrypted at both the transport and message levels.

- Transport level encryption is provided by HTTPS.
- Messages will be sent/received via the AS2 specification and will be signed/encrypted using digital certificate-based authentication.

TGA specific validation rules

M.1 ICH ICSR message header specifications

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
M.1.1	messagetype	Yes		All ICSRs must be 'ichicsr', including reports from clinical trials.	If validation fails for any of M.1 data elements, the error message returned will be: <DTD Descriptor>: 'The incoming message has failed batch validation. Please refer to the TGA EDI ICSR R2 Implementation Guidelines'.
M.1.2	messageformatversion	No			
M.1.3	messageformatrelease	No			
M.1.4	messagenumb	Yes			
M.1.5	messagesenderidentifier	Yes			
M.1.6	messagereceiveridentifier	Yes			
M.1.7a	messagedateformat	Yes			
M.1.7b	messagedate	Yes	The date specified cannot refer to a future date.		

ICH ICSR M2 data processing specifications

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
	safetyreportversion	Yes	5. The version number cannot exceed 2 numerics. 6. The version number must be greater than any previous versions of the same ICSR (A.1.0.1) that has been sent unless the ICSR is a nullification or the transmission date (A.1.3b) is greater than any previous versions.		Default error message. If validation fails, the following error message is returned: <safetyreportid>: 'Invalid data supplied. The ICSR version number has been previously submitted.'

A. E2B (R2) ICSR specifications

A.1 Safety report

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules and notes	Notes	Acknowledgement response error messages
A.1.0.1	safetyreportid	Yes	1. The safetyreportid cannot exceed 100 alphanumeric characters. 2. The safetyreportid value must start with a valid ISO 2 character country code. 3. A batch cannot contain multiple ICSRs with the same safetyreportid. 4. The worldwide unique case identification number must not change between versions of the same ICSR (safetyreportid). 5. The safetyreportid must not have been previously nullified.	1. Where a safetyreportid supplied in a batch, exceeds its length, the batch will be rejected. 2. The value should be the country code (Same as at A.1.1)-company or regulator name-report number. For example, a case from Australia would use “AU-companyname-12345”, where 12345 is a company’s unique case report number.	1. If validation fails for any of M.1 data elements, the error message returned will be: <safetyreportid>: 'The incoming message has failed batch validation. Please refer to the TGA EDI ICSR R2 Implementation Guidelines'. 2. Default error message. The following error message is returned: <safetyreportid>: ‘Invalid data supplied. The batch contains two or more ICSR messages with the same Sender's Case Safety Report Unique Identifier.’ 3. The following error message is returned: <safetyreportid>: ‘Invalid data supplied. The worldwide unique case identification number has changed from the previous version.’

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules and notes	Notes	Acknowledgement response error messages
					4. The following error message is returned: <safetyreportid>: 'Invalid data supplied. The ICSR has been nullified'
A.1.1	primarysourcecountry	No		This should correspond to one of the primary source countries reported in A.2.1.3.	Default error message.
A.1.2	occurcountry	No		Include if different from A.1.1	Default error message.
A.1.3a	transmissiondateformat	Yes			Default error message.
A.1.3b	transmissiondate	Yes	The date specified cannot refer to a future date.		Default error message.
A.1.4	reporttype	Yes	If A.2.3.3 is provided against the 1st primary source record, this field must be provided with a value of '2'.		Default error message.
A.1.5.1	serious	Yes	If any of the A.1.5.2 fields are set to 1 (Yes), then A.1.5.1 must be '1' (Yes).	If any of the reactions or events in the case is serious, the serious flag is to be set to Yes.	Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules and notes	Notes	Acknowledgement response error messages
A.1.5.2	seriousnessdeath	Yes	If A.1.5.1 is '1' (yes), at least one of the A.1.5.2 fields must be '1' (yes). The A.1.5.2 data elements which are not set to '1' (yes) can either be set to '2' (no) or left as null/absent.		Default error message.
	seriousnesslifethreatening	Yes		Life threatening is defined in the ICH E2A	
	seriousnesshospitalization	Yes			
	seriousnessdisabling	Yes			
	seriousnesscongenitalanomali	Yes			
	seriousnessother	Yes		Other means medically important condition as defined in ICH E2A	
A.1.6a	receivedateformat	Yes			Default error message.
A.1.6b	receivedate	Yes	The date specified cannot refer to a future date.		Default error message.
A.1.7a	receiptdateformat	Yes			Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules and notes	Notes	Acknowledgement response error messages
A.1.7b	receiptdate	Yes	The date specified cannot refer to a future date.		Default error message.
A.1.8.1	additionaldocument	Conditional	If A.1.8.2 is provided, A.1.8.1 must be '1' (Yes).		Default error message.
A.1.8.2	documentlist	Conditional	A.1.8.2 must be provided if A.1.8.1 is '1' (Yes).		Default error message.
A.1.9	fulfillexpeditecriteria	No			Default error message.
A.1.10.1	authoritynumb	Conditional	Either A.10.1 **or** A.1.10.2 must be provided.		Default error message.
A.1.10.2	companynumb	Conditional			
A.1.11	duplicate	Conditional	If A.1.11.1 and A.1.11.2 are provided, A.1.11 must be provided with a value of '1' (Yes).		Default error message.
A.1.11.1	duplicatesource	Conditional	If A.1.11 is '1' (Yes), A.1.11.1 must be provided. If A.1.11.2 is provided, A.1.11.1 must be provided.	Details from previous transmissions of the same case from either other senders or by the same sender using a different reporting channel such as email or online form should be provided in this section.	Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules and notes	Notes	Acknowledgement response error messages
A.1.11.2	duplicatenumb	Conditional	If A.1.11 is '1' (Yes), A.1.11.2 must be provided. If A.1.11.1 is provided, A.1.11.2 must be provided.		Default error message.
A.1.12	linkreportnumb	No		This field identifies reports or cases that warrant being evaluated together. For example, a sequence of cases involving the same patient.	Default error message.
A.1.13	casenullification	Conditional	If provided, must be '1' (Yes). If A.1.13.1 is provided, A.1.13 must be provided with value '1' (Yes).	This section indicates that a previous report should be considered void (nullified). To resubmit reports that have previously been nullified, assign a new safetyreportid.	Default error message.
A.1.13.1	nullificationreason	Conditional	When A.1.13 is '1' (Yes), A.1.13.1 must be provided		Default error message.
A.1.14	medicallyconfirm	No			Default error message.

A.2 Primary source(s) of information

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
A.2	primarysource	Yes	At least one of the data elements in section A.2.1 must be supplied in the message (excluding A.2.1.1 elements).		Default error message.
A.2.1.1a	reportertitle	No	If provided, this element will be stored by the TGA as Masked.	You do not need to provide this element.	Default error message.
A.2.1.1.b	reportergivename	No	If provided, this element will be stored by the TGA as Masked.	You do not need to provide this element.	Default error message.
A.2.1.1c	reportermiddlename	No	If provided, this element will be stored by the TGA as Masked.	You do not need to provide this element.	Default error message.
A.2.1.1d	reporterfamilyname	No	If provided, this element will be stored by the TGA as Masked.	You do not need to provide this element.	Default error message.
A.2.1.2a	reporterorganization	Conditional			Default error message.
A.2.1.2b	reporterdepartment	Conditional			Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
A.2.1.2c	reporterstreet	Conditional	If A.2.1.4 does not equal '5', this element will be stored by the TGA as Masked.	The TGA does not collect this element for consumers and non-health professionals.	Default error message.
A.2.1.2d	reportercity	Conditional	If A.2.1.4 does not equal '5', this element will be stored by the TGA as Masked.	The TGA does not collect this element for consumers and non-health professionals.	Default error message.
A.2.1.2e	reporterstate	Conditional			Default error message.
A.2.1.2f	reporterpostcode	Conditional	If A.2.1.4 does not equal '5', this element will be stored by the TGA as Masked.	The TGA does not collect this element for consumers and non-health professionals.	Default error message.
A.2.1.3	reportercountry	Conditional			Default error message.
A.2.1.4	qualification	Conditional			Default error message.
A.2.2	Literaturereference	No		References are to be provided in the Vancouver Convention.	Default error message.
A.2.3.1	studyname	No			Default error message.
A.2.3.2	sponsorstudynumb	No			Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
A.2.3.3	observestudytype	Conditional	If A.1.4 is'2', A.2.3.3 must be provided for the 1st primary source record (Report from Study).		Default error message.

A.3 Information on sender and receiver of ISCR

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
A.3.1.1	sendertype	No		Sender Type should be provided but when it is not provided it will default to unknown.	Default error message.
A.3.1.2	senderorganization	Yes			Default error message.
A.3.1.3a	senderdepartment	No			Default error message.
A.3.1.3b	sendertitle	No	If provided, this element will be stored by the TGA as Masked.	You do not need to provide this element.	Default error message.
A.3.1.3c	sendergivename	No	If provided, this element will be stored by the TGA as Masked.	You do not need to provide this element.	Default error message.
A.3.1.3d	sendermiddlename	No	If provided, this element will be stored by the TGA as Masked.	You do not need to provide this element.	Default error message.
A.3.1.3e	senderfamilyname	No	If provided, this element will be stored by the TGA as Masked.	You do not need to provide this element.	Default error message.
A.3.1.4a	senderstreetaddress	No			Default error message.
A.3.1.4b	sendercity	No			Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
A.3.1.4c	senderstate	No			Default error message.
A.3.1.4d	senderpostcode	No			Default error message.
A.3.1.4e	sendercountrycode	No			Default error message.
A.3.1.4f	sendertel	No			Default error message.
A.3.1.4g	sendertelextension	No			Default error message.
A.3.1.4h	sendertelcountrycode	No			Default error message.
A.3.1.4i	senderfax	No			Default error message.
A.3.1.4j	senderfaxextension	No			Default error message.
A.3.1.4k	senderfaxcountrycode	No			Default error message.
A.3.1.4l	senderemailaddress	No		An email address should be provided. This email address may be used when sending correspondence to Sender.	Default error message.
A.3.2.1	receivertype	No		Where provided it should be set to a value of '2' Regulatory Authority	Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
A.3.2.2a	receiverorganization	No		Where provided it should be set to 'TGA'	Default error message.
A.3.2.2b	receiverdepartment	No		Where provided it should be set to 'Pharmacovigilance and Special Access Branch'	Default error message.
A.3.2.2c	receivertitle	No			Default error message.
A.3.2.2d	receivergivename	No			Default error message.
A.3.2.2e	receivermiddlename	No			Default error message.
A.3.2.2f	receiverfamilyname	No			Default error message.
A.3.2.3a	receiverstreetaddress	No		Where provided it should be set to 'PO Box 100'	Default error message.
A.3.2.3b	receivercity	No		Where provided it should be set to 'Woden'	Default error message.
A.3.2.3c	receiverstate	No		Where provided it should be set to 'ACT'	Default error message.
A.3.2.3d	receiverpostcode	No		Where provided it should be set to '2606'	Default error message.
A.3.2.3e	receivercountrycode	No		Where provided it should be set to 'AU'	Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
A.3.2.3f	receivertel	No			Default error message.
A.3.2.3g	receivertelextension	No		Where provided it should be set to '262216963'	Default error message.
A.3.2.3h	receivertelcountrycode	No		Where provided it should be set to '+61'	Default error message.
A.3.2.3i	receiverfax	No			Default error message.
A.3.2.3j	receiverfaxextension	No		Where provided it should be set to '262031713'	Default error message.
A.3.2.3k	receiverfaxcountrycode	No		Where provided it should be set to '+61'	Default error message.
A.3.2.3l	receiveremailaddress	No		Where provided it should be set to 'adr.reports@tga.gov.au'	Default error message.

B. Information on the case

B.1 Patient characteristics

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1	patient	Yes	At least one of the following must be provided: · Patient initials (B.1.1); or · A patient record number (B.1.1.1.a – d); or · Patient age information – B.1.2.1a/b (DOB), B.1.2.2a/b (Age at time of onset), B.1.2.2.1a/b (gestational age); or · B.1.2.3 (Age group); or · Patient sex (B.1.5).		Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.1	patientinitial	Conditional		Where the data is known but cannot be provided this element can be set to PRIVACY or MSK. Where you believe there is a real patient involved but do not have any specific patient characteristics available, this element can be set to UNK.	Default error message.
B.1.1.1a	patientgpmedicalrecordnumb	Conditional			Default error message.
B.1.1.1b	patientspecialistrecordnumb	Conditional			Default error message.
B.1.1.1c	patienthospitalrecordnumb	Conditional			Default error message.
B.1.1.1d	patientinvestigationnumb	Conditional			Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.2	Age information			Only one element describing age should be provided, based on the most precise information available.	Default error message.
B.1.2.1a	patientdatebirthformat	Conditional	If B.1.2.1b is provided, B.1.2.1a b must be provided.		Default error message.
B.1.2.1b	patientdatebirth	Conditional	Where B.1.2.1a is provided, B.1.2.1b must be provided.		Default error message.
B.1.2.2a	patientonsetage	Conditional	Where B.1.2.2a is provided, B.1.2.2b must be provided. Age must be less than or equal to 120 years.	If several reactions or events are in the case, the age at the time of the first reaction or event should be used.	Default error message.
B.1.2.2b	patientonsetageunit	Conditional	Where B.1.2.2b is provided, B.1.2.2a must be provided.		Default error message.
B.1.2.2.1a	gestationperiod	Conditional	Where B.1.2.2.1a is provided, B.1.2.2.1b must be provided.		Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.2.2.1b	gestationperiodunit	Conditional	Where B.1.2.2.1b is provided, B.1.2.2.1a must be provided.		Default error message.
B.1.2.3	patientagegroup	Conditional			Default error message.
B.1.3	patientweight	No	Weight must be less than or equal to 500kg. Up to 4 decimal places are allowed and this is included in the character limit.		Default error message.
B.1.4	patientheight	No	Height must be less than or equal to 300 cm.		Default error message.
B.1.5	patientsex	Conditional		If sex is unknown, intersex or indeterminate leave this data element blank.	Default error message.
B.1.6a	lastmenstrualdateformat	Conditional	Where B.1.6b is provided, B.1.6a must be provided.		Default error message

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.6b	patientlastmenstrualdate	Conditional	Where B.1.6a is provided, B.1.6b must be provided. The date specified cannot refer to a future date.		Default error message.

B.1.7 Relevant medical history and concurrent conditions

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.7.1a.1	patientepisodenamemeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.
B.1.7.1a.2	patientepisodename	No	Must match either an 8 digit MedDRA code or text for a Low Level Term.	Where possible, use the 8 digit MedDRA code for a Low Level Term rather than the text for the LLT.	Default error message.
B.1.7.1b	patientmedicalstartdateformat	Conditional	If B.1.7.1c is provided, B.1.7.1b must be provided.		Default error message.
B.1.7.1c	patientmedicalstartdate	Conditional	Where B.1.7.1b is provided, B.1.7.1c must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.7.1d	patientmedicalcontinue	No			Default error message.
B.1.7.1e	patientmedicalenddateformat	Conditional	Where B.1.7.1f is provided, B.1.7.1e must be provided.		Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.7.1f	patientmedicalenddate	Conditional	Where B.1.7.1e is provided, B.1.7.1f must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.7.1g	patientmedicalcomment	No			Default error message.
B.1.7.2	patientmedicalhistorytext	No			Default error message.

B.1.8 Relevant past drug history

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.8a	patientdrugname	Conditional	If any of the following data elements in the patient block (B.1.8b+B.1.8c or B.1.8d+B.1.8e or B.1.8f.2 or B.1.8fg.2) are provided, B.1.8a must be provided.	Where possible provide the trade name of the product.	Default error message.
B.1.8b	patientdrugstartdateformat	Conditional	Where B.1.8c is provided, B.1.8 b must be provided.		Default error message.
B.1.8c	patientdrugstartdate	Conditional	Where B.1.8b is provided, B.1.8c must be provided.		Default error message.
B.1.8d	patientdrugenddateformat	Conditional	Where B.1.8e provided, B.1.8d must be provided.		Default error message.
B.1.8e	patientdrugenddate	Conditional	Where B.1.8d provided, B.1.8e must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.8f.1	patientindicationmeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.8f.2	patientdrugindication	No	Must match either an 8 digit MedDRA code or text for a Low Level Term.	Where possible, use the 8 digit MedDRA code for a Low Level Term rather than the text for the LLT.	Default error message.
B.1.8g.1	patientdrgreactionmeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.
B.1.8g.2	patientdrgreaction	No	Must match either an 8 digit MedDRA code or text for a Low Level Term.	Where possible, use the 8 digit MedDRA code for a Low Level Term rather than the text for the LLT.	Default error message.
B.1.9	patientdeath	No			Default error message.
B.1.9.1a	patientdeathdateformat	Conditional	Where B.1.9.1b is provided, B.1.9.1a must be provided.		Default error message.
B.1.9.1b	patientdeathdate	Conditional	Where B.1.9.1a is provided, B.1.9.1b must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.9.2.a	patientdeathreportmeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.9.2.b	patientdeathreport	No	Must match either an 8 digit MedDRA code or text for a Low Level Term.	Where possible, use the 8 digit MedDRA code for a Low Level Term rather than the text for the LLT.	Default error message.
B.1.9.3	patientautopsyyesno	Conditional	If B.1.9.1b is provided, B.1.9.3 must be provided. If B.1.9.4b is provided, B.1.9.3must be '1' (Yes).		Default error message.
B.1.9.4a	patientdetermautopsmeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.
B.1.9.4b	patientdetermineautopsy	Conditional	Where B.1.9.4b provided, B.1.9.3 must be = '1' (Yes).		Default error message.

B.1.10 For a parent-child/foetus report, Information concerning the parent

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.10.1	parentidentification	No			Default error message.
B.1.10.2.1a	parentbirthdateformat	Conditional	Where B.1.10.2.1b is provided, B.1.10.2.1a must be provided.	Only one element describing age should be used, based on the most precise information available.	Default error message.
B.1.10.2.1b	parentbirthdate	Conditional	If B.1.10.2.1a provided, B.1.10.2.1b must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.10.2.2a	parentage	Conditional	If B.1.10.2.2a is provided, B.1.10.2.2b must be supplied. Age must be less than or equal to 120 years.		Default error message.
B.1.10.2.2b	parentageunit	Conditional	If B.1.10.2.2b is provided, B.1.10.2.2a must be supplied.		Default error message.
B.1.10.3a	parentlastmenstrualdateformat	Conditional	Where B.1.10.3b is provided, B.1.10.3a must be provided.		Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.10.3b	parentlastmenstrualdate	Conditional	Where B.1.10.3a provided, B.1.10.3b must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.10.4	parentweight	No	Weight must be less than or equal to 500kg. Up to 3 decimal places are allowed and this is included in the character limit.		Default error message.
B.1.10.5	parentheight	No	Where provided height must be less than or equal to 300 cm.		Default error message.
B.1.10.6	parentsex	No			Default error message.
B.1.10.7.1a.1	parentmdepisodemeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.
B.1.10.7.1a.2	parentmedicalepisodename	No	Must match either an 8 digit MedDRA code or text for a Low Level Term.	Where possible, use the 8 digit MedDRA code for a Low Level Term rather than the text for the LLT.	Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.10.7.1b	parentmedicalstartdateformat	Conditional	Where B.1.10.7.1c is provided, B.1.10.7.1b must be provided.		Default error message.
B.1.10.7.1c	parentmedicalstartdate	Conditional	Where B.1.10.7.1b is provided, B.1.10.7.1c must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.10.7.1d	parentmedicalcontinue	No			Default error message.
B.1.10.7.1e	parentmedicalenddateformat	Conditional	Where B.1.10.7.1f is provided, B.1.10.7.1e must be provided.		Default error message.
B.1.10.7.1f	parentmedicalenddate	Conditional	Where B.1.10.7.1e is provided, B.1.10.7.1f must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.10.7.1g	parentmedicalcomment	No			Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.10.7.2	parentmedicalrelevanttext	No			Default error message.
B.1.10.8a	parentdrugname	Conditional	If any of the following fields in the parent block (B.1.10.8b and B.1.10.8c or B.1.10.8d and B.1.10.8e or B.1.10.8f.2 or B.1.10.8g.2) are provided, B.1.10.8a must be provided	Where possible provide the trade name of the product.	Default error message.
B.1.10.8b	parentdrugstartdateformat	Conditional	If B.1.10.8c provided, B.1.10.8b must be provided.		Default error message.
B.1.10.8c	parentdrugstartdate	Conditional	Where B.1.10.8b provided, B.1.10.8c must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.10.8d	parentdrugenddateformat	Conditional	Where B.1.10.8e is provided, B.1.10.8d must be provided.		Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.1.10.8e	parentdrugenddate	Conditional	Where B.1.10.8d is provided, B.1.10.8e must be provided. The date specified cannot refer to a future date.		Default error message.
B.1.10.8f.1	parentdrgindicationmeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.
B.1.10.8f.2	parentdrugindication	No	Must match either an 8 digit MedDRA code or text for a Low Level Term.	Where possible, use the 8 digit MedDRA code for a Low Level Term rather than the text for the LLT.	Default error message.
B.1.10.8g.1	parentdrgreactionmeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.
B.1.10.8g.2	parentdrugreaction	No	Must match either an 8 digit MedDRA code or text for a Low Level Term.	Where possible, use the 8 digit MedDRA code for a Low Level Term rather than the text for the LLT.	Default error message.

B.2 Reaction(s) or event(s)

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.2.i.0	primarysourcereaction	No			Default error message.
B.2.i.1.a	reactionmeddraversionllt	Yes	A decimal point/full stop is allowed. For example, 19.1.	Only one MedDRA version may be used in ICSR's received through the EDI. The first instance of the MedDRA version populated in B.2.i.1.a will be used for validation of all other MedDRA version data elements in the ICSR.	Default error message.
B.2.i.1.b	reactionmeddrallt	Yes	Must match either an 8 digit MedDRA code or text for a Low Level Term.	Where possible, use the 8 digit MedDRA code for a Low Level Term rather than the text for the LLT.	Default error message.
B.2.i.2.a	reactionmeddraversionpt	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.
B.2.i.2.b	reactionmeddrapt	No			Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.2.i.3	termhighlighted	No		If the information is not explicitly provided by the initial reporter, the term should not be considered a highlighted term. The seriousness of the reaction or event should be based on the ICH E2A criteria.	Default error message.
B.2.i.4a	reactionstartdateformat	Conditional	Where B.2.i.4b is provided, B.2.i.4a must be provided.		Default error message.
B.2.i.4b	reactionstartdate	Conditional	Where B.2.i.4a is provided, B.2.i.4b must be provided. The date specified cannot refer to a future date.		Default error message.
B.2.i.5a	reactionenddateformat	Conditional	Where B.2.i.5b is provided, B.2.i.5a must be provided.		Default error message.
B.2.i.5b	reactionenddate	Conditional	Where B.2.i.5a is provided, B.2.i.5b must be provided. The date specified cannot refer to a future date.		Default error message.
B.2.i.6a	reactionduration	Conditional	Where B.2.i.6a is provided, B.2.i.6b must be provided.		Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.2.i.6b	reactiondurationunit	Conditional	Where B.2.i.6b is provided, B.2.i.6a must be provided.		Default error message.
B.2.i.7.1a	reactionfirsttime	Conditional	Where B.2.i.7.1a is provided, B.2.i.7.1b must be provided.		Default error message.
B.2.i.7.1b	reactionfirsttimeunit	Conditional	Where B.2.i.7.1b is provided, B.2.i.7.1a must be provided.		Default error message.
B.2.i.7.2a	reactionlasttime	Conditional	Where B.2.i.7.2a is provided, B.2.i.7.2b must be provided.		Default error message.
B.2.i.7.2b	reactionlasttimeunit	Conditional	Where B.2.i.7.2b is provided, B.2.i.7.2a must be provided.		Default error message.
B.2.i.8	reactionoutcome	No			Default error message.

B.3 Results of tests and procedures relevant to the investigation of the patient

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.3.1a	testdateformat	Conditional	Where B.3.1b is provided, B.3.1a must be provided.		Default error message.
B.3.1b	testdate	Conditional	If B.3.1a provided, B.3.1b must be provided. The date specified cannot refer to a future date.		Default error message.
B.3.1c	testname	Conditional	If B.3.1b provided, B.3.1c must be provided.		Default error message.
B.3.1d	testresult	No		Test results should be reported where known.	Default error message.
B.3.1e	testunit	No			Default error message.
B.3.1.1	lowtestrange	No			Default error message.
B.3.1.2	hightestrange	No			Default error message.
B.3.1.3	moreinformation	No			Default error message.
B.3.2	resultstestsprocedures	No			Default error message.

B.4 Drug(s) information

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.4.k.1	drugcharacterization	Yes	Each ICSR must contain at least one 'Suspect' or 'Interacting' drug.	Where 'Interacting' has been set for a product in the ICSR, information should be provided on the other product/s that were part of the interaction.	Default error message.
B.4.k.2.1	medicinalproduct	Conditional	Either B.4.k.2.1 or B.4.k.2.2 must be provided.	Where possible provide the trade name of the product.	Default error message.
B.4.k.2.2	activesubstancename	Conditional	Either B.4.k.2.1 or B.4.k.2.2 must be provided.		Default error message.
B.4.k.2.3	obtaindrugcountry	No			Default error message.
B.4.k.3	drugbatchnumb	No			Default error message.
B.4.k.4.1	drugauthorizationnumb	No		Where possible provide the Australian Register of Therapeutic Goods (ARTG) number of the product.	Default error message.
B.4.k.4.2	drugauthorizationcountry	Conditional	Where B.4.k.4.1 is provided, B.4.k.4.2 is required.	Set this field to 'AU' when providing the ARTG number in B.4.k.4.1.	Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.4.k.4.3	drugauthorizationholder	No			Default error message.
B.4.k.5.1	drugstructuredosagenumb	Conditional	Where B.4.k.5.2 is provided, B.4.k.5.1 must be provided.		Default error message.
B.4.k.5.2	drugstructuredosageunit	Conditional	Where B.4.k.5.1 is provided, B.4.k.5.2 must be provided.		Default error message.
B.4.k.5.3	drugseparatedosagenumb	No			Default error message.
B.4.k.5.4	drugintervaldosageunitnumb	Conditional	Where B.4.k.5.4 is provided, B.4.k.5.5 must be provided.		Default error message.
B.4.k.5.5	drugintervaldosagedefinition	Conditional	Where B.4.k.5.5 is provided, B.4.k.5.4 must be provided.		Default error message.
B.4.k.5.6	drugcumulativedosagenumb	Conditional	Where B.4.k.5.6 is provided, B.4.k.5.7 must be provided.		Default error message.
B.4.k.5.7	drugcumulativedosageunit	Conditional	Where B.4.k.5.7 is provided, B.4.k.5.6 must be provided.		Default error message.
B.4.k.6	drugdosagetext	No			Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.4.k.7	drugdosageform	No			Default error message.
B.4.k.8	drugadministrationroute	No			Default error message.
B.4.k.9	drugparadministration	No			Default error message.
B.4.k.10a	reactiongestationperiod	Conditional	Where B.4.k.10a is provided, B.4.k.10b must be provided.		Default error message.
B.4.k.10b	reactiongestationperiodunit	Conditional	Where B.4.k.10b is provided, B.4.k.10a must be provided.		Default error message.
B.4.k.11a	drugindicationmeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.
B.4.k.11b	drugindication	No		If there are multiple indications for use in the case for the product, enter the main indication in B.4.k.11b and include information on the additional indications within the case narrative (B.5.1). Alternatively, the sender can provide more than one iteration (k) for the same drug.	Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.4.k.12a	drugstartdateformat	Conditional	Where B.4.k.12b is provided, B.4.k.12a must be provided.		Default error message.
B.4.k.12b	drugstartdate	Conditional	Where B.4.k.12a is provided, B.4.k.12b must be provided. The date specified cannot refer to a future date.		Default error message.
B.4.k.13.1a	drugstartperiod	Conditional	Where B.4.k.13.1a is provided, B.4.k.13.1b must be provided.	B.4.k.13 captures the interval between each drug and only the reaction or event in the first iteration of B.2.i.	Default error message.
B.4.k.13.1b	drugstartperiodunit	Conditional	Where B.4.k.13.1b is provided, B.4.k.13.1a must be provided.		Default error message.
B.4.k.13.2a	druglastperiod	Conditional	Where B.4.k.13.2a is provided, B.4.k.13.2b must be provided.		Default error message.
B.4.k.13.2b	druglastperiodunit	Conditional	Where B.4.k.13.2b is provided, B.4.k.13.2a must be provided.		Default error message.
B.4.k.14a	drugenddateformat	Conditional	Where B.4.k.14b provided, B.4.k.14a must be provided.		Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.4.k.14b	drugenddate	Conditional	Where B.4.k.14a provided, B.4.k.14b must be provided. The date specified cannot refer to a future date.	Leave blank for ongoing drug administration following onset of reaction or event	Default error message.
B.4.k.15a	drugtreatmentduration	Conditional	Where B.4.k.15a is provided, B.4.k.15b must be provided.		Default error message.
B.4.k.15b	drugtreatmentdurationunit	Conditional	Where B.4.k.15b is provided, B.4.k.15a must be provided.		Default error message.
B.4.k.16	actiondrug	No		'Not applicable' should be used in circumstances such as if the patient died or the treatment had been completed prior to the reaction or event.	Default error message.
B.4.k.17.1	drugrecurreadministration	No		'Unknown' indicates that a rechallenge was done, but it is unknown if the event recurred. This data element should not be completed if it is unknown whether a rechallenge was done.	Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.4.k.17.2a	drugrecuractionmeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a.		Default error message.
B.4.k.17.2b	drugrecuraction	No			Default error message.
B.4.k.18.1a	drugreactionassesmeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a.	Generally ordered from the most important or the most serious to the least important.	Default error message.
B.4.k.18.1b	drugreactionasses	No	Must be the same as a provided reaction/event in MedDRA terminology (LLT) specified in B.2.i.1.b or the same as the MedDRA preferred term (PT) specified in B.2.i.2.b.	Where drugreactionasses is provided, the source, result and method of assessment should be completed.	Default error message.
B.4.k.18.2	drugassessmentsource	No			Default error message.
B.4.k.18.3	drugassessmentmethod	No			Default error message.
B.4.k.18.4	drugresult	No			Default error message.

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.4.k.19	drugadditional	No		B.4.k.19 may be used to specify additional information pertinent to the case that is not covered by above sections (for example, beyond expiration date, batch and lot tested and found to be within specifications) or additional information on the indication for the drug.	Default error message.

B.5 Narrative case summary and further information

E2B R2 element	DTD descriptor	Mandatory field	TGA specific validation rules	Notes	Acknowledgement response error messages
B.5.1	narrativeincludeclinical	No		Focused, factual, and clear description of the case should be given, including the words or short phrases used by the reporter.	Default error message.
B.5.2	reportercomment	No		Reporter's comments on the diagnosis, causality assessment, or other issues considered relevant.	Default error message.
B.5.3a	senderdiagnosismeddraversion	Conditional	Must be the same as the MedDRA version at B.2.i.1.a		Default error message.
B.5.3b	senderdiagnosis	No		Opportunity to combine signs and symptoms that were reported into a succinct diagnosis.	Default error message.
B.5.4	sendercomment	No		Information concerning the sender's assessment of the case and can be used to describe disagreement with, and/or alternatives to the diagnoses given by the initial reporter.	Default error message.

Test scenarios to use the E2B standard

The 10 test cases below should be submitted using the TGA validation rules and those in the ICH ISCR DTD as part of the registration process for E2B.

TGA is aware that not all marketing authorisation holders will be running post marketing studies of clinical trials, however all tests need to be carried out. If necessary you can use fictional information so that if you do start running these studies, you won't need to carry out testing again.

Tests

1. Create an ICSR including details of other drugs, medical history, drug history, narrative (B.5.1) and tests, showing:
   - both the structured medical (B.1.7.1a-1g and B.1.7.2), and drug (B.1.8a – B.1.8g.2) history
   - examples of correctly structured tests (B.3.1a - B.3.1.3 and B.3.2) and show the pharmaceutical forms (B.4.k.7) have been incorporated correctly.
2. Create a fatal ICSR with cause of death and post mortem details where:
   - seriousness death flag must be yes
   - reaction outcome must be fatal
   - patient death date should be provided (B.1.9.1(a+b))
   - reported cause of death should be provided (B.1.9.2(a+b))
   - autopsy flag (B.1.9.3) should be set to 'yes'
   - autopsy cause of death should be provided (B.1.9.4 (a+b)).
3. A nullification ICSR:
   - the nullification flag must be set to Yes (A.1.13)
   - the nullification reason (A.1.13.1) must be provided.
4. A parent-child ICSR where:
   - the child/foetus must be encoded as the patient
   - the parent's details must be provided in the parent's section
   - it is essential that:
     - § the child (patient) drug route of administration (B.4.k.8) must be transmammary, transplacental, or other (if parent is male)
     - § the parent route of administration is entered (B.4.k.9)
     - § data element B.1.10 contains validating parent details (initials, age, sex, weight) as well as any relevant parent history and other details available

5. A follow-up ICSR to an initial ICSR which you have previously sent:
   - initial report should have the same receive (A.1.6(a+b)) and receipt (A.1.7(a+b)) dates
   - follow-up report must have the same receive date as the initial but the receipt date must be changed to reflect when the follow-up information was received
   - the sender's case safety report unique ID (A.1.0.1) and worldwide case ID (A.1.10) must be identical in both the initial and follow-up reports.
6. A follow-up ICSR to an initial ICSR which has previously been sent from an alternative sender (for example, alternative company or competent authority) should be:
   - the worldwide case ID (A.1.10) must be the same as was sent by the previous sender of the ICSR
   - the sender's case safety report unique ID (A.1.0.1) will differ
7. An ICSR from a study where:
   - report type must be 'Report from study' (A.1.4)
   - study type must be 'Other studies' (A.2.3.3)
   - study name and number should be provided (A.2.3.1 and A.2.3.2).
8. An ICSR based on a literature article where the literature reference (A.2.2) must be provided in Vancouver style.
9. An ICSR with duplicate details completed (A.1.11.1 and A.1.11.2) to capture:
   - all the safety report IDs that have been used in the past to transmit a case between different organisations
   - all previous paper numbers that have been used in the past, to enable the detection of duplicates if a case had previously been sent on paper (for example, ADRS Case number) and company CIOMS MFR Control numbers.
10. A suspected unexpected serious adverse reaction (SUSAR) where the:
    - report type must be 'Report from study' (A.1.4)
    - study type must be 'Clinical trials' (A2.3.3)
    - study name must be provided (A.2.3.1)
    - study number must be provided (A2.3.2)
    - patient ID must be provided (B.1.1.1d).

Version history

Version	Description of change	Author	Effective date
V1.0	Original publication	Technical and Safety Improvement Section	June 2018
V1.1	Minor update to correct errors	Technical and Safety Improvement Section	July 2018
V1.2	Minor update to reflect system changes	Technical and Safety Improvement Section	December 2018
V1.3	Updated email address	Technical and Safety Improvement Section	October 2019

Therapeutic Goods Administration

PO Box 100 Woden ACT 2606 Australia
Email: info@tga.gov.au Phone: 1800 020 653 Fax: 02 6232 8605
http://www.tga.gov.au

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