

Annex I

List of the names, pharmaceutical forms, strength of the veterinary medicinal products, animal species, routes of administration, marketing authorisation holders in the Member States

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Austria	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol 25 mg/ml Lösung zum Eingeben über das Trinkwasser für Hühner	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Austria	Bayer Animal Health 51368 Leverkusen, Germany	Baycox Direct 25 mg/ml Lösung zum Eingeben über das Trinkwasser für Hühner und Puten	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Belgium	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 25mg/ml, solution pour administration dans l'eau de boisson pour poulets et dindes / oplossing voor gebruik in drinkwater voor kippen en kalkoenen toltrazuril / Lösung zum Eingeben über das Trinkwasser für Hühner und Puten	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Belgium	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 2,5% 25 mg/ml, solution buvable / drank / Lösung zum Einnehmen	Toltrazuril	25 mg/ml	Oral solution	Chicken, turkey	Oral use
Belgium	Avimedical B.V. Abbinkdijk 1, 7255 LX Hengelo Gld The Netherlands	Coccinox	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Belgium	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Bulgaria	Pharma Vet Ltd Shumen, ul. "Otec Paisij" № 40, Bulgaria	Фармакокс разтвор Pharmacox liquid	Toltrazuril	2.5 g/ 100 ml	Oral solution	Chicken, turkey	Oral use
Bulgaria	Interchemie Werken De Adelaar BV Metaalweg 8, 5804 CG Venray, The Netherlands	Intracox oral	Toltrazuril	25 mg/ml	Oral solution	Chicken, turkey	Oral use
Bulgaria	Kepro B.V. Maagdenburgstraat 17, 7421 ZA Deventer The Netherlands	Кепкокс 2.5% орал /Kepcox 2.5% oral	Toltrazuril	25 mg/ml	Oral solution	Chicken, turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Bulgaria	Ceva Santé Animale 10, Av. de La Ballastière 33500 Libourne France	Севазурил/Cevazuril	Toltrazuril	25 mg/ml	Oral solution	Broilers, growing parents of chickens, bright, turkeys	Oral use
Bulgaria	Bayer Animal Health 51368 Leverkusen, Germany	Байкокс 2,5% разтвор/Baycox 2,5% solution	Toltrazuril	25 mg/ml	Oral solution	Chicken, turkey	Oral use
Bulgaria	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril 25 mg/ml разтвор за перорална употреба с питейната вода при пилета / Dozuril 25 mg/ml solution for use in drinking water for chicken	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Croatia	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 25 mg/ml, otopina za primjenu u vodi za piće, za kokoši i purane	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Cyprus	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 2,5% w/v Πόσιμο διάλυμα για ορνίθια κρεοπαραγωγής και ινδοορνίθια.	Toltrazuril	25 mg/ml	Oral solution	Broiler, turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Czech Republic	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 2,5% (w/v) perorální roztok	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broiler), turkey	Oral use
Denmark	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Estonia	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Estonia	Interchemie Werken De Adelaar Eesti AS Vanapere tee 14, Puunsi Village, Viimsi Rural Municipality, Harju County, 74013 Estonia	Intracox Oral	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey	Oral use
Estonia	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
France	Ceva Santé Animale 10, Av. de La Ballastière 33500 Libourne France	Cevazuril 25 mg/ml solution buvable pour poulets de chair, poulettes et reproducteurs	Toltrazuril	25 mg/ml	Oral solution	Chicken (broilers, pullets, breeders)	Oral use
France	Virbac 1ere avenue, 2065 m, LID, 06516 Carros, France	Toltracox 2,5%	Toltrazuril	25 mg/ml	Oral solution	Chicken (broilers, pullets, breeders)	Oral use
France	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol 25 mg/ml solution buvable pour poulets	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
France	VetPharma Animal Health S.L. Les Corts 23, 08028 Barcelona, Spain	Zorabel 25 mg/ml solution pour administration dans l'eau de boisson pour poulets et dindes	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
France	Elanco GmbH Heinz-Lohmann-Straße 4 27472 Cuxhaven Germany	Baycox 25 mg/ml solution pour administration dans l'eau de boisson des poules et des dindes	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
France	Avimedical B.V. Abbinkdijk 1, 7255 LX Hengelo Gld The Netherlands	Coccinox 25 mg/ml solution orale pour administration dans l'eau de boisson pour poulets et dindes	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey, pigeon	Oral use
France	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Cozuril CT 25 mg/ml solution pour administration dans l'eau de boisson pour poulets et dindes	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Germany	VetPharma Animal Health S.L. Les Corts 23, 08028 Barcelona, Spain	Zorabel 25 mg/ml Lösung zum Eingeben über das Trinkwasser für Hühner und Puten	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Germany	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril CT	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Germany	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol 25 mg/ml Lösung zum Eingeben über das Trinkwasser	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Germany	Elanco GmbH Heinz-Lohmann-Straße 4 27472 Cuxhaven Germany	Baycox 25 mg/ml, Lösung zum Eingeben über das Trinkwasser für Hühner und Puten	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Germany	Laboratorios Karizoo, S.A. Mas Pujades, 11-12, Pol. Ind. La Borda, Caldes de Montbui, 08140 Barcelona Spain	Toltra-K 25 mg/ml Lösung zum Eingeben über das Trinkwasser für Hühner und Puten	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Germany	Bremer Pharma GmbH, Werkstr. 42, D-34414 Warburg, Scherfede, Germany	Bremacox 25 mg/ml Lösung zum Eingeben über das Trinkwasser für Hühner und Puten	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Germany	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril 25 mg/ml Lösung zum Eingeben über das Trinkwasser für Hühner	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Greece	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 2,5%	Toltrazuril	25 mg/ml	Oral solution	Broiler, turkey	Oral use
Greece	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Greece	VetPharma Animal Health S.L. Les Corts 23, 08028 Barcelona, Spain	Zorabel	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Greece	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol, Διάλυμα για χορήγηση με πόσιμο νερό 25 mg/ml	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Hungary	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 2,5% koncentrátum belsőleges oldathoz A.U.V.	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Hungary	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril 25 mg/ml, oldat ivónízbe keveréshez háziyúkoknak A.U.V.	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Hungary	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol 25 mg/ml oldat ivónízbe keveréshez háziyúkok részére A.U.V.	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Hungary	Laboratorios Karizoo, S.A. Mas Pujades, 11-12, Pol. Ind. La Borda, Caldes de Montbui, 08140 Barcelona Spain	Toltra-K 25 mg/ml oldat ivóvízbe keveréshez háziyúkok és pulykák számára A.U.V.	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Ireland	Elanco GmbH Heinz-Lohmann-Straße 4 27472 Cuxhaven Germany	Baycox 25 mg/ml, solution for use in drinking water for chickens and turkeys	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey	Oral use
Ireland	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril 25 mg/ml solution for use in drinking water for chickens	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Ireland	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol 25 mg/ml solution for use in drinking water for chickens	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Italy	Elanco Italia S.p.A Via dei Colatori, 12 50019 Sesto Fiorentino (FI) Italy	Baycox PT 25 mg/ml soluzione per uso in acqua di bevanda per polli da carne e tacchini	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Italy	Elanco Italia S.p.A Via dei Colatori, 12 50019 Sesto Fiorentino (FI) Italy	Baycox Soluzione orale 25 mg/ml per polli da carne e tacchini	Toltrazuril	25 mg/ml	Oral solution	Broiler, turkey	Oral use
Italy	Ceva Salute Animale S.p.A. Viale Colleoni 15 20864 Agrate Brianza (MB) Italy	Cevazuril, 25 mg/ml, soluzione orale, per polli da carne e tacchini, da somministrare diluita in acqua da bere.	Toltrazuril	25 mg/ml	Oral solution	Broiler, turkey	Oral use
Italy	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril CT 25 mg/ml, soluzione per l'uso in acqua da bere per polli e tacchini	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Italy	Laboratorios Karizoo, S.A. Mas Pujades, 11-12, Pol. Ind. La Borda, Caldes de Montbui, 08140 Barcelona Spain	Toltra-K-25 Soluzione da usare nell'acqua di bevanda per polli e tacchini	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Italy	Industria Italiana Integratori Trei S.p.A. Viale Corassori, 62 41100 Modena Italy	Treizuril, 25 mg/ml, soluzione orale per polli da carne e tacchini	Toltrazuril	25 mg/ml	Oral solution	Broiler, turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Italy	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol 25 mg/ml soluzione per uso in acqua da bere per polli	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Latvia	Interchemie Werken De Adelaar Eesti AS Vanapere tee 14, Puunsi Village, Viimsi Rural Municipality, Harju County, 74013 Estonia	LV Intracox Oral 25 mg/ml šķīdums iekšķīgai lietošanai tītariem un vistām	Toltrazuril	25 mg/ml	Oral solution	Chicken, turkey	Oral use
Latvia	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril 25 mg/ml šķīdums lietošanai ar dzeramo ūdeni vistām	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Latvia	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol 25 mg/ml šķīdums lietošanai ar dzeramo ūdeni vistām	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Lithuania	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril 25 mg/ml, tirpalas naudoti su geriamuoju vandeniu vištoms	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Lithuania	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol 25 mg/ml, tirpalas, naudoti su geriamuoju vandeniu vištoms	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Lithuania	UAB Interchemie werken De Adelaar LT, Terminalo str. 13, Biruliškes, 54469, Kaunas distr. Lithuania	Intracox oral, 25 mg/ml tirpalas naudoti su ger. vandeniu vištoms ir kalakutams	Toltrazuril	25 mg/ml	Oral solution	Chicken (broiler), turkey	Oral use
Luxembourg	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 25 mg/m, solution pour administration dans l'eau de boisson pour poulets et dindes / Lösung zum Eingeben über das Trinkwasser für Hühner und Puten	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Luxembourg	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 2,5% 25 mg/ml, solution buvable / Lösung zum Einnehmen	Toltrazuril	25 mg/ml	Oral solution	Chicken, turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Luxembourg	Avimedical B.V. Abbinkdijk 1, 7255 LX Hengelo Gld The Netherlands	Coccinox	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Luxembourg	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Malta	Kepro B.V. Maagdenburgstraat 17, 7421 ZA Deventer The Netherlands	Kepcox 2.5%	Toltrazuril	25 mg/ml	Oral solution	Chicken (pullets, breeders), turkey	Oral use
Netherlands	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 25 mg/ml, oplossing voor gebruik in drinkwater voor kippen en kalkoenen	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey	Oral use
Netherlands	Avimedical B.V. Abbinkdijk 1, 7255 LX Hengelo Gld The Netherlands	Coccinox 25 mg/ml orale oplossing voor gebruik in drinkwater voor kippen, kalkoenen, duiven en konijnen	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey, pigeon	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Netherlands	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril CT 25 mg/ml oplossing voor toediening in drinkwater voor kippen en kalkoenen	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Netherlands	Pantex Holland B.V. Smaragdweg 15 5525 LA Hapert The Netherlands	Toltrapan 25 mg/ml, oplossing voor gebruik in drinkwater voor kippen en kalkoenen	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey	Oral use
Netherlands	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril 25 mg/ml oplossing voor toediening in drinkwater voor kippen	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Poland	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 2,5%, (2,5 g/100 ml, roztwór doustny dla kur i indyków)	Toltrazuril	2.5 g/ 100 ml	Oral solution	Chicken (pullets), turkey	Oral use
Poland	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Poland	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril CT	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey	Oral use
Poland	Biofaktor Sp. z o. o. Czysta 4 st., 96-100 Skierniewice, Poland	Endocox	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Poland	Lovapharm Consulting B.V. Rijsven 3 5645 KH Eindhoven The Netherlands	Lovacox oral	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, pullets, breeders), turkey	Oral use
Poland	Laboratorios Karizoo, S.A. Mas Pujades, 11-12, Pol. Ind. La Borda, Caldes de Montbui, 08140 Barcelona Spain	Toltra-K	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Poland	VetPharma Animal Health S.L. Les Corts 23, 08028 Barcelona, Spain	Zorabel	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Poland	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Portugal	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 25 mg/ml, solução para administração na água de bebida para galinhas e perus	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Portugal	VetPharma Animal Health S.L. Les Corts 23, 08028 Barcelona, Spain	Zorabel 25 mg/ml solução para administração na água de bebida para galinhas e perus	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
Portugal	Laboratorios Karizoo, S.A. Mas Pujades, 11-12, Pol. Ind. La Borda, Caldes de Montbui, 08140 Barcelona Spain	Toltra-K 25 mg/ml solução para administração na água de bebida para galinhas e perus	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Portugal	Calier Portugal, S.A. Centro Empresarial Sintra- Estoril II, Ed. C, R. Pé de Mouro Estrada de Albarraque 2710 - 335 Sintra Portugal	Zuritol 25 mg/ml solução para administração na água de bebida para galinhas	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Romania	Ceva Santé Animale 10, Av. de La Ballastière 33500 Libourne France	Cevazuril	Toltrazuril	25 mg/ml	Oral solution	Broilers, replaceme nts, pullets, turkeys	Oral use
Romania	Dopharma Research B.V. Zalmweg 24 4941 VX Raamsdonksveer The Netherlands	Dozuril 25 mg/ml	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use
Romania	Bayer Animal Health 51368 Leverkusen, Germany	Baycox	Toltrazuril	25 mg/ml	Oral solution	Chicken, turkey	Oral use
Slovak Republic	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 2,5% (w/v) perorálny roztok	Toltrazuril	25 mg/ml	Oral solution	Chicken, turkey	Oral use
Slovenia	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 25 mg/ml peroralna raztopina za piščance in purane	Toltrazuril	25 mg/ml	Oral solution	Chicken, turkey	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
Spain	Bayer Animal Health 51368 Leverkusen, Germany	Baycox 25 mg/ml solución para administración en agua de bebida para aves	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (broilers, chick for replaceme nt), turkey	Oral use
Spain	Laboratorios Karizoo, S.A. Mas Pujades, 11-12, Pol. Ind. La Borda, Caldes de Montbui, 08140 Barcelona Spain	Toltra-K 25 mg/ml solución para administración en agua de bebida para pollos y pavos	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Spain	VetPharma Animal Health S.L. Les Corts 23, 08028 Barcelona, Spain	Zorabel 25 mg/ml solucion para administracion en agua de bebida para pollos y pavos	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders), turkey	Oral use
Spain	Laboratorios Calier, S.A. Barcelonès, 26 (Pla del Ramassà) 08520 Les Franqueses del Vallès (Barcelona) Spain	Zuritol 25 mg/ml solucion para uso en agua de bebida para pollos	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken (pullets, breeders)	Oral use

Member State EU/EEA	Marketing authorisation holders	Name	INN	Strength	Pharmaceutical form	Animal species	Route of administration
UK (Northern Ireland) ¹	Elanco Europe Ltd, Form 2, Bartley Way, Bartley Wood Business Park, Hook, RG 27 9XA, United Kingdom	Baycox 25 mg/ml, solution for use in drinking water for chickens and turkeys	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
UK (Northern Ireland) ¹	Laboratorios Karizoo, S.A. Mas Pujades, 11-12, Pol. Ind. La Borda, Caldes de Montbui, 08140 Barcelona Spain	Toltra-K 25 mg/ml solution for use in drinking water for chickens and turkeys	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use
UK (Northern Ireland) ¹	VetPharma Animal Health S.L. Les Corts 23, 08028 Barcelona, Spain	Zorabel 25 mg/ml solution for use in drinking water for chickens and turkeys	Toltrazuril	25 mg/ml	Solution for use in drinking water	Chicken, turkey	Oral use

¹For the United Kingdom, as from 1 January 2021, EU Law applies only to the territory of Northern Ireland (NI) to the extent foreseen in the Protocol on Ireland/NI.

Annex II

Scientific conclusions and grounds for amendment of the summary of product characteristics

Overall summary of the scientific evaluation of veterinary medicinal products containing toltrazuril to be administered orally to chickens (see Annex I)

1. Introduction

Toltrazuril is an anticoccidial of the triazinetrione group and it is active against *Eimeria* spp. Veterinary medicinal products containing toltrazuril are used in chickens, amongst other animal species, for the treatment of coccidiosis, which is a serious disease for chickens, especially affecting the young chicks.

Veterinary medicinal products containing toltrazuril are not authorised for use in chickens producing eggs for human consumption. However, in exceptional cases, young layers may also become infected by particular *Eimeria* species. If no authorised alternative treatment is available, toltrazuril may be used in layers in accordance with Article 113 of Regulation (EU) 2019/6 (the so-called "cascade") and a withdrawal period of at least 10 days must be applied to eggs in those cases (in line with Article 115 (1)(c) of Regulation (EU) 2019/6).

In 2020, an incident was reported in EudraVigilance Veterinary concerning the use of a toltrazuril product in the Netherlands in young layers of approximately 20 weeks of age to treat an infection with *Eimeria brunetti* causing illness and mortality. The recommended dose of 7 mg of toltrazuril per kg bodyweight per day for 2 days was applied in the drinking water and a withdrawal period of 7 days was advised by the attending veterinarian (in line with Article 11 of Directive 2001/82/EC, which was in force at that time). Nevertheless, the Dutch Inspection Unit requested a check for residues (the marker residue being toltrazuril sulfone, also known as ponazuril) before eggs could enter the food chain. The results demonstrated that the marker residue could be identified in treated layers up to 96 days after treatment.

Due to the long retention of toltrazuril in eggs from treated birds, the Netherlands considered that the restriction period of 4 weeks before the onset of lay (as indicated in the product information of a number of veterinary medicinal products authorised in the EU) might not be sufficient to ensure compliance with the recommended MRL and might be too short to ensure that residues in a standard food basket including eggs would not exceed the acceptable daily intake, and therefore, there might be a potential risk for the consumer.

In view of the above, the Netherlands considered it necessary to refer the matter to the CVMP in the interest of protecting consumer safety in the European Union.

Therefore, on 10 December 2021, the Netherlands initiated a procedure under Article 35 of Directive 2001/82/EC, for veterinary medicinal products containing toltrazuril to be administered orally to chickens. The Committee for Veterinary Medicinal Products was requested to review all available residue data and recommend an appropriate restriction period before the onset of lay during which the above-mentioned veterinary medicinal products must not be used in chickens.

Although at the start of this referral procedure, a maximum residue limit (MRL) for toltrazuril in chicken eggs was not formally established in the European Union (EU), it is important to note that pursuant to Article 9(1)(b) of Regulation (EC) No 470/2009, the Netherlands submitted to the European Medicines Agency on 29 June 2021 a request for the extension of maximum residue limits for toltrazuril to chicken eggs. On 9 December 2021, the CVMP issued an opinion recommending the establishment of an MRL for the marker residue toltrazuril sulfone of 140 µg/kg for poultry eggs².

² EMA. Summary of opinion on the extension of maximum residue limits for toltrazuril to chicken eggs (EMA/CVMP/709072/2021) – [link](#)

2. Discussion of data available

Qualitative and quantitative composition

The Committee noted that veterinary medicinal products concerned by this referral procedure have a similar composition. Given that the veterinary medicinal products are for oral use and that all authorised generics are bioequivalent to the reference product ('Baycox 25 mg/ml'; Marketing Authorisation Holder (MAH): Bayer Animal Health GmbH/Elanco Animal Health Inc.), the CVMP considered that any differences in the composition are not expected to influence the restriction period in eggs.

Therefore, the CVMP considered that all residue data for all concerned veterinary medicinal products can be considered together for the determination of a suitable period before the onset of lay in which the products must not be used. Consequently, the Committee concluded that the outcome of the assessment regarding the consumer safety can be applied to all products concerned by this referral procedure.

Information related to eggs as stated in section 4.11 (QRD template v8.2) of the summary of product characteristics for the different veterinary medicinal products

The Committee noted that in most of the veterinary medicinal products concerned by this referral the summary of product characteristics (SPC) contained statements clarifying that they are not authorised for use in laying birds. Moreover, several SPCs indicated that the veterinary medicinal product should not be used within 4 weeks of onset of laying. However, the Committee noted that no information was provided on how the length of this restriction period was determined.

Residue data

No proprietary data relating to residue kinetics of toltrazuril in chicken eggs following oral administration were provided by the MAHs. However, published literature (three publications from scientific journals provided by some MAHs concerned by this referral) were made available to the CVMP: Mulder *et al.*, 2005³; Varenina *et al.*, 2012⁴; Pietruk *et al.*, 2021⁵. In addition, an unpublished residue depletion study was made available by the European Reference Laboratory for residues in Berlin⁶ and some residue data were obtained from EudraVigilance Veterinary in 2020.

The data covered various ranges of sampling times but only the EURL Technical report⁶ had a relevant range of sampling times, since it was the only study in which chickens were treated at approximately 4 weeks before the onset of lay. This time of treatment is more relevant as compared to the other studies in which laying birds were used, because it followed the restriction period that is indicated in the product information of several veterinary medicinal products authorised in the European Union (i.e., 4 weeks before the start of the laying period), which constitutes the scope of this referral procedure. Therefore, the CVMP considered the EURL study⁶ as the pivotal study for the determination of the restriction period before the onset of lay in which birds must not be treated.

In the EURL study⁶, 10 healthy brown laying hens (15 weeks of age, bodyweight: 1.04-1.22 kg) were given the veterinary medicinal product 'Toltra K' in the drinking water at a dose of 7 mg of toltrazuril per kg bodyweight per day for 2 days. Eggs were collected daily for approximately 5 weeks from the

³ Mulder P.P.J., *et al.* (2005). Deposition and depletion of the coccidiostats toltrazuril and halofuginone in eggs. *Analytica chimica acta*, 529(1-2), 331-337.

⁴ Varenina I., *et al.* (2012). Deposition of toltrazuril residues in eggs during and after treatment with dose over the therapeutic dose. [Conference poster]. Board of the EuroResidue Conferences Foundation. Residues of veterinary drugs in food. Proceedings of the EuroResidue VII Conference, Egmond aan Zee, The Netherlands, 14-16 May 2012. Volume 1, 2 and 3 pp. 533-538.

⁵ Pietruk K., *et al.* (2021). Risk of residues of toltrazuril sulfone in eggs after oral administration – could setting maximum residue limit be helpful?. *Food chemistry*, 360, 130054.

⁶ Technical Report 2019/2020 of the EURL for residues (EURL Berlin, 2020)

onset of lay and were analysed for the concentration of the marker residue toltrazuril sulfone using a liquid chromatography with tandem mass spectrometry detection (LC-MS/MS) method with a decision limit (CCa) of 4.42 µg/kg for toltrazuril sulfone, 6.47 µg/kg for toltrazuril and 0.49 µg/kg for toltrazuril sulfoxide.

Toltrazuril sulfone residues were found in eggs up to 8 weeks after treatment. The individual residue concentrations declined from 167.49 µg of toltrazuril sulfone per kg at day 27 after treatment, to below CCa (4.42 µg of toltrazuril sulfone per kg) at day 55 after treatment. The residue concentrations of toltrazuril and toltrazuril sulfoxide were below their respective CCa at all time points.

The residue levels of toltrazuril sulfone were below the CVMP recommended MRL value for eggs (*i.e.*, 140 µg/kg) at day 28 after treatment. There was only one sample above the recommended MRL, which corresponded to the first egg collected (on day 27 after treatment) and the concentration was 167.49 µg of toltrazuril sulfone per kg. On the first sampling time points only one egg per day was present (not all birds started laying at the same time) and from the later time points, there is a substantial variation in residue concentrations which had to be taken into consideration. Log-linear regression revealed that the intercept of the regression line with the recommended MRL is at 30 days and the 95/95 tolerance limit reaches the recommended MRL at 38 days.

The CVMP considered that the most suitable statistical approach is the "SCPM method" which is a method used for the determination of the withdrawal period for milk (EMEA/CVMP/473/98-FINAL)⁷ in which the 95/95 tolerance limit for each time point is compared to the MRL. For this method, it is not necessary to know which eggs were laid by which hen and the method assumes a log-normal distribution of the residues at each time point.

According to this analysis, the restriction period between the administration and the start of the laying period would be 41 days, which is also in line with the MAHs' proposal that suggested to set the restriction period at 6 weeks, based on the data from Pietruk *et al.* (2021)⁵ in which the 95/95 tolerance limit would also be between 35-40 days. The Committee also noted that the EudraVigilance Veterinary data are consistent with this proposal since the residues were below the recommended MRL from day 39 after treatment onwards.

Overall, from the residue datasets, the Committee considered that the restriction period of 4 weeks before the onset of lay is not sufficient to ensure the consumer safety according to the CVMP recommended MRL value for poultry eggs (140 µg/kg). It is important to note that although at the time of concluding this referral procedure this MRL has not yet been formally adopted and included in Table 1 of the Annex to Regulation (EU) No. 37/2010, it does not change the fact that the CVMP considers it to represent a level of residues that can be considered safe, from a consumer protection perspective. As the recommended MRL is considered a safe value, it has been used as a reference point in this referral procedure for the derivation of an appropriate time interval between administration and the onset of lay.

Therefore, the CVMP considered that a restriction period of 6 weeks before the onset of lay, in which birds must not be treated, would ensure that residues of the marker residue toltrazuril sulfone in eggs are below the recommended MRL of 140 µg/kg from the time the birds start laying, and hence, it is considered adequate to protect consumers. Therefore, the CVMP considered that the product information for all concerned veterinary medicinal products should be amended accordingly to reflect this outcome.

⁷ CVMP Note for guidance for the determination of withdrawal periods for milk (EMEA/CVMP/473/98-FINAL) - [link](#)

3. Benefit-risk assessment

Introduction

The CVMP was requested to review all available residue data in chicken eggs for the veterinary medicinal products containing toltrazuril to be administered orally to chickens and to recommend an appropriate restriction period before the onset of lay, during which chickens must not be treated with the above-mentioned veterinary medicinal products.

Benefit assessment

While the efficacy of the concerned veterinary medicinal products in chickens has not been specifically assessed as part of this referral procedure, the products under assessment are considered to be effective in chickens for the treatment of coccidiosis which is a serious disease, especially affecting the young chicks.

Risk assessment

Quality, target animal safety, user safety and the environmental risk for the concerned veterinary medicinal products have not been assessed in this referral procedure. Furthermore, for generic products, bioequivalence was not evaluated, since this was already done within the respective marketing authorisation procedures.

Toltrazuril was previously evaluated by the CVMP in order to establish maximum residue limits (MRLs). On 9 December 2021, the CVMP adopted an opinion recommending the extension of MRLs for toltrazuril to chicken eggs². Furthermore, with reference to Article 5 of Regulation (EC) No 470/2009 and in line with the criteria laid down in Commission Regulation (EU) 2017/880, the Committee agreed to extrapolate the conclusions to poultry eggs (CVMP recommended MRL for toltrazuril in poultry eggs: 140 µg/kg).

Coccidiosis mostly occurs in young chicks (most often between 4-6 weeks of age), but it may occasionally occur at a somewhat later age. Because toltrazuril sulfone, the main metabolite, has a relatively long elimination half-life, residues of this metabolite may be excreted in eggs, even if the administration took place at a timepoint before the onset of laying. Therefore, a restriction period of 4 weeks before the onset of lay in which birds must not be treated is currently included in the product information of some veterinary medicinal products concerned by this referral procedure.

However, a risk has been identified regarding the length of the above-mentioned authorised restriction period for chickens. The available information indicated that it may be insufficient to allow residues in eggs to fall below the recommended MRL (i.e., 140 µg/kg), thereby posing a risk to consumers.

Risk management or mitigation measures

It should be noted that the residues in eggs from turkeys are not within the scope of this referral and therefore, have not been assessed within this procedure.

On the basis of residue data in chickens, a restriction period of 6 weeks before the onset of lay, in which birds must not be treated have been derived for all veterinary medicinal products concerned by this referral procedure. This period ensures that residues of the marker residue toltrazuril sulfone in eggs are below the CVMP recommended MRL of 140 µg/kg from the time the birds start laying, and hence, it is considered adequate to ensure consumer safety.

Evaluation and conclusions on the benefit-risk balance

Having considered the grounds for this referral procedure and the data available, the CVMP concluded that the restriction period before the onset of lay, during which chickens must not be treated with

orally administered veterinary medicinal products containing toltrazuril should be amended as recommended.

The overall benefit-risk balance for the veterinary medicinal products containing toltrazuril to be administered orally to chickens remains positive, subject to the recommended changes in the product information.

Grounds for amendment of the summary of product characteristics, labelling and package leaflet

Whereas

- on the basis of the available residue depletion data, the CVMP considered that the restriction period before the onset of lay during which orally administered veterinary medicinal products containing toltrazuril must not be used in chickens should be amended to provide assurance for consumer safety;
- the CVMP considered that the overall benefit-risk balance for the products under this procedure remains positive subject to amendments in the product information

the CVMP has recommended the amendment of the marketing authorisations for veterinary medicinal products containing toltrazuril to be administered orally to chickens as referred in Annex I for which the summary of product characteristics, labelling and package leaflet are set out in Annex III.

Annex III

Amendments in the relevant sections of the Summary of product characteristics, labelling and package leaflet

Summary of product characteristics

4.11 Withdrawal period(s) (QRD template v8.2) / **3.12 Withdrawal periods** (QRD template v9.0)

...

Not for use in birds producing or intended to produce eggs for human consumption. Do not use within 6 weeks before the start of the laying period.

Labelling

8. WITHDRAWAL PERIOD(S) (QRD template v8.2) / 7. WITHDRAWAL PERIODS (QRD template v9.0)

...

Not for use in birds producing or intended to produce eggs for human consumption. Do not use within 6 weeks before the start of the laying period.

Package leaflet

10. WITHDRAWAL PERIOD(S) (QRD template v8.2) / **10. Withdrawal periods** (QRD template v9.0)

...

Not for use in birds producing or intended to produce eggs for human consumption. Do not use within 6 weeks before the start of the laying period.