Table 1: General recommendations for systemic drugs for AE adult patients, who are candidates for systemic treatment (for details see corresponding chapter)

	Conve	ntional systemic t	reatments	Biologics			Rescue therapy		
	Ciclosporin	Methotrexate	Azathioprine	Dupilumab	Tralokinumab	Abrocitinib	Baricitinib	Upadacitinib	Systemic corticosteroids
Recommendation	$\uparrow \uparrow$	1	1	ተተ	ተተ	$\uparrow \uparrow$	ተተ	ተተ	1
Dose for adults <sup>1</sup>	licensed ≥ 16 years; standard dosage adults: 2.5-5 mg/kg per day in two single doses	off-label; commonly used dosage adults: initial dose: 5-15 mg/ per week; maximum dose: 25 mg/ week	off-label; commonly used dosage adults: 1-3 mg/kg per day	licensed ≥ 6 months; adults: initially 600 mg s.c. day 1 followed by 300 mg Q2W	licensed for adults; initially 600 mg s.c. day 1 followed by 300 mg Q2W; consider Q4W dosing at week 16 in those achieving clear or almost clear skin	licensed for adults; dosage adults: 200 mg per day, reduction to 100 mg per day possible, depending on treatment response; age ≥ 65: 100 mg per day; the lowest effective dose for maintenance should be considered	licensed for adults; dosage adults: 4 mg per day, reduction to 2 mg per day possible, depending on treatment response	licensed ≥ 12 years; dosage adults: 15 or 30 mg per day based on individual patient presentation; age ≥ 65: 15 mg per day; the lowest effective dose for maintenance should be considered	general unspecific licence for adults and children for steroid responsive skin disease; dosage maximum: 1 mg/kg per day
Time to response (weeks) <sup>2</sup>	1-2	8-12	8-12	4-6	4-8	1-2	1-2	1-2	1-2
Time to relapse (weeks, based on expert experience) <sup>2</sup>	<2	>12	>12	>8	> 8	<2	<2	<2	<2
Monitoring	complete blood count, renal and liver profile, blood pressure,	complete blood count, renal and liver profile, PIIINP if available, screen for chronic infections	complete blood count, renal and liver profile, TPMT activity if available, screen for chronic infections	not required	not required	complete blood count, lipid profile, liver profile	complete blood count, lipid profile, liver profile	complete blood count, lipid profile, liver profile	not required for short-term treatment, consider blood glucose and testing for adrenal gland suppression with high doses/longer- term treatment

Selection of most	serum	nausea,	gastrointestinal	Conjunctivitis,	upper	upper respiratory tract	upper respiratory	upper	skin atrophy,
relevant adverse	creatinine $\uparrow$ ,	fatigue,	disturbances,	upper	respiratory	infections,,	tract infections,,	respiratory tract	weight gain,
events	blood	liver enzymes	idiosyncratic	respiratory	tract	increase in LDL	increase in LDL	infections, acne;	sleep
	pressure $\uparrow$	个,	hypersensitivity	tract	infections;	cholesterol;	cholesterol;	headache,	disturbance,
		myelotoxicity	reactions,	infections,	conjunctitivitis	thrombocytopenia,	thrombocytosis,	anaemia and	mood changes,
			hepatotoxicity,	arthralgia		increased creatine	nausea and	neutropenia, CK	hyperglycaemia
			myelotoxicity			phosphokinase,	abdominal pain	elevation,	or new onset
						nausea and abdominal	herpes virus	increase in LDL	diabetes,
						pain	infections,	cholesterol,	peptic
						herpes virus	acne	nausea and	ulcers/gastritis,
						infections,		abdominal pain	osteoporosis
						acne		herpes virus	
								infections	

<sup>1</sup>SmPC, <sup>2</sup>expert experience, ↑ rise, AE- atopic eczema; GL – guideline, LDL – low density lipoprotein, PIIINP - Procollagen III N-Terminal Propeptide, TPMT – Thiopurine-S-Methyltransferase

Symbols	Implications (adapted from GRADE <sup>1</sup> )				
<u>^</u>	/e believe that all or almost all informed people would make that choice.				
1	e believe that most informed people would make that choice, but a substantial number would not.				
0	We cannot make a recommendation.				
$\checkmark$	We believe that most informed people would make a choice against that intervention, but a substantial number would not.				
$\downarrow\downarrow$	We believe that all or almost all informed people would make a choice against that choice.				
	No recommendation.				

	Conventional systemic treatments		Biologics			Rescue therapy			
	Ciclosporin	Methotrexate	Azathioprine	Dupilumab	Tralokinumab	Abrocitinib	Baricitinib	Upadacitinib	Systemic corticosteroids
Children and adolescents with AE who are candidates for systemic treatment	<b>↑</b> ↑	Ť	Ŷ	<b>↑</b> ↑	<b>↑</b> ↑	<b>↑</b> ↑		ተተ	
Dose for children	licensed for ≥ 16 years commonly used dosage children: 2.5-5 mg/kg per day in two single doses	off-label; commonly used dosage children: 0.3– 0.4 mg/kg per week	off label; commonly used dosage children: 1-3 mg/kg per day	licensed for $\geq$ 6 months; age 6 months-6 years: from 5kg <15 kg 200 mg Q4W, 15kg <30 kg 300 mg Q4W age 6-11: from 15kg <60kg, initially 300 mg s.c. day 1 &15 followed by 300 mg Q4W, when $\geq$ 60 kg, initially 600 mg s.c. day 1 followed by 300 mg Q2W age 12-17: <60 kg: initially 400 mg s.c. day 1 followed by 200 mg Q2W, when $\geq$ 60 kg: initially 600 mg s.c. day 1 followed by 300 mg Q2W	licensed for ≥ 12 years; initially 600 mg s.c. day 1 followed by 300 mg Q2W; consider Q4W dosing at week 16 in those achieving clear or almost clear skin	off-label; only in UK approved ≥ 12 years; commonly used dosage children: 100 mg per day	off-label	licensed for ≥ 12 years; age 12-17 (>= 30 kg bw): 15 mg per day	general unspecific licence for children for steroid responsive skin disease;; dosage maximum: 1 mg/kg per day
<b>Pregnancy</b> (in candidates for systemic treatment)	Ŷ	$\downarrow\downarrow$	Ŷ	0	0	$\downarrow\downarrow$	$\downarrow\downarrow$	$\downarrow\downarrow$	↑ prednisolone (0.5mg/kg/d) o <i>nly</i> as rescue therapy for acute flares
Breastfeeding	Ŷ	Ŷ	¥	0	0	Ŷ	Ŷ	Ŷ	↑ prednisolone (0.5mg/kg/d) o <i>nly</i> as rescue therapy for acute flares

## Table 2: General recommendations for systemic drugs for special AE patient populations (for details see corresponding chapter)

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<sup>1</sup>SmPC; Q2W - once every 2 weeks

Sy	mbols	Implications (adapted from GRADE <sup>1</sup> )			
	ተተ	Ne believe that all or almost all informed people would make that choice.			
	1	e believe that most informed people would make that choice, but a substantial number would not.			
	0	We cannot make a recommendation.			
	$\checkmark$	We believe that most informed people would make a choice against that intervention, but a substantial number would not.			
	$\uparrow \uparrow$	We believe that all or almost all informed people would make a choice against that choice.			
		No recommendation.			

## Table 3: General recommendations for topical drugs for treatment of atopic eczema (for details see corresponding chapter)

Overall recommendation	TCS	$\uparrow \uparrow$	TCI↑↑			
	TCS class I and II	TCS class III and IV	Tacrolimus 0.1% Tacrolimus 0.03%	Pimecrolimus 1%		
For further information see background text	class I not suitable for long-term proactive treatment; long-term proactive treatment only class II	acute flare; proactive treatment with TCS class III class IV <i>not</i> for long term daily treatment or head and neck; class IV not recommended for proactive treatment either	acute flare; long-term proactive treatment; especially in face, intertriginous sites, anogenital area	acute flare; especially in face, intertriginous sites, anogenital area		
Most important side effects	skin atrophy telangiectasia striae distensae ecchymosis hypertrichosis perioral dermatitis	skin atrophy telangiectasia striae distensae ecchymosis hypertrichosis perioral dermatitis corticosteroid addiction syndrome suppression of adrenal function	initial warmth, tingling or burning	initial warmth, tingling or burning		
	TCI class II and III are off lab		in label for proactive treatment	not suitable for proactive treatment		
Special considerations						
Suitable for <b>children &gt; 2 to &lt; 16</b> years of age	yes	yes	yes (0.03%) <sup>2</sup>	yes <sup>2</sup>		
Suitable for <b>babies</b> < 2 years of age	yes	under specialist supervision	yes (0.03%) <sup>1</sup>	yes <sup>2</sup> (from the age of three months)		
Suitable during <b>pregnancy</b>	yes	yes	yes (0.03% & 0.1%) <sup>1</sup>	yes <sup>1</sup>		
Suitable during <b>breastfeeding</b>	yes	yes	yes (0.03% & 0.1%) <sup>1</sup>	yes <sup>1</sup>		
Suitable for pruritus	yes	yes	yes (0.03% & 0.1%)	yes		

## <sup>1</sup> off label use <sup>2</sup> licensed use

Symbols	Implications (adapted from GRADE <sup>1</sup> )			
<u>ተተ</u>	Ne believe that all or almost all informed people would make that choice.			
$\uparrow$	/e believe that most informed people would make that choice, but a substantial number would not.			
0	We cannot make a recommendation.			
$\checkmark$	We believe that most informed people would make a choice against that intervention, but a substantial number would not.			
$\downarrow\downarrow$	We believe that all or almost all informed people would make a choice against that choice.			
	No recommendation.			

## **References**

[1] Kaminski-Hartenthaler A, Meerpohl JJ, Gartlehner G, Kien C, Langer G, Wipplinger J, et al. [GRADE guidelines: 14. Going from evidence to recommendations: the significance and presentation of recommendations]. Z Evid Fortbild Qual Gesundhwes. 2014;108; 413-420.