# Annex III

## Amendments to relevant sections of the Product Information

*Note:*

These amendments to the relevant sections of the Product Information are the outcome of the referral procedure.

The product information may be subsequently updated by the Member State competent authorities, in liaison with the Reference Member State, as appropriate, in accordance with the procedures laid down in Chapter 4 of Title III of Directive 2001/83/EC.

## Amendments to relevant sections of the Product Information

The existing product information shall be amended (insertion, replacement or deletion of the text as appropriate) to reflect the agreed wording as provided below.

### A. Summary of Product Characteristics

- **Section 4.2 Posology and method of administration**

*This following wording should be reflected in this section. Text in strikethrough should be deleted from the summary of product characteristics, if present.*

**250 mg tablets**

Dosage is determined by the intensity of the pain or fever and individual sensitivity of response to <invented name>. It is essential to choose the lowest dose that controls pain and fever.

For children and adolescents up to 14 years of age 8–16 mg metamizole per kg body weight can be given as a single dose. In case of fever, a dose of 10 mg metamizole per kilogram body weight is generally sufficient for children. Adults and adolescents of 15 years of age or older (> 53 kg) can take up to 1,000 mg metamizole as a single dose, which can be taken up to 4 times daily at intervals of 6–8 hours corresponding to a maximum daily dose of 4,000mg.

A clear effect can be expected 30 to 60 minutes after oral administration.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | tablets | mg | tablets | mg |
| 32 - 53 | 10-14 years | 1 - 2 | 250 – 500 | 8 | 2,000 |
| >53 | ≥ 15 years | 2 - 4 | 500-1,000 | 16 | 4,000 |

*Paediatric population*

<invented name> is not recommended for children younger than 10 years of age due to the fixed amount of 250 mg metamizole contained in one tablet. Other pharmaceutical forms/strengths are available which can be appropriately dosed for smaller children.

**500 mg tablets/capsules (also applicable for 500mg powder for oral solution/effervescent powder/granules for oral solution)**

Dosage is determined by the intensity of the pain or fever and individual sensitivity of response to <invented name>. It is essential to choose the lowest dose that controls pain and fever.

Adults and adolescents of 15 years of age or older (> 53 kg) can take up to 1,000 mg metamizole as a single dose, which can be taken up to 4 times daily at intervals of 6–8 hours corresponding to a maximum daily dose of 4,000mg.

A clear effect can be expected 30 to 60 minutes after oral administration.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | tablets /sachets | mg | tablets/sachets | mg |
| >53 | ≥ 15 years | 1-2 | 500-1,000 | 8 | 4,000 |

*Paediatric population*

<invented name> is not recommended for children younger than 15 years of age due to the fixed amount of 500mg metamizole contained in one tablet/capsule/sachet. Other pharmaceutical forms/strengths are available which can be appropriately dosed for smaller children.

**575 mg capsules**

Dosage is determined by the intensity of the pain or fever and individual sensitivity of response to <invented name>. It is essential to choose the lowest dose that controls pain and fever.

Adults and adolescents of 15 years of age or older (> 53 kg) can take 575 mg metamizole as a single dose, which can be taken up to 6 times daily at intervals of 4 - 6 hours corresponding to a maximum daily dose of 3,450mg.

A clear effect can be expected 30 to 60 minutes after oral administration.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | capsules | mg | capsules | mg |
| >53 | ≥ 15 years | 1 | 575 | 6 | 3,450 |

*Paediatric population*

<invented name> is not recommended for children younger than 15 years of age due to the fixed amount of 575mg metamizole contained in one capsule. Other pharmaceutical forms/strengths are available which can be appropriately be dosed for smaller children.

**500 mg/ml oral drops, solution (1 ml = 20 drops)**

Dosage is determined by the intensity of the pain or fever and individual sensitivity of response to <invented name>. It is essential to choose the lowest dose that controls pain and fever.

In children and adolescents up to 14 years old, 8–16 mg metamizole per kg body weight can be given as a single dose. In the case of fever, a dose of 10 mg metamizole per kilogram body weight is generally sufficient for children. Adults and adolescents from 15 years of age (>53 kg) can take up to 1,000 mg as a single dose. In dependence on the daily maximum dose a single dose can be taken up to 4 times daily at intervals of 6–8 hours.

A clear effect can be expected 30 to 60 minutes after oral administration.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | drops | mg | drops | mg |
| <9 | <12 months | 1-5 | 25-125 | 4-20 | 100-500 |
| 9-15 | 1-3 years | 3-10 | 75-250 | 12-40 | 300-1,000 |
| 16-23 | 4-6 years | 5-15 | 125-375 | 20-60 | 500-1,500 |
| 24-30 | 7-9 years | 8-20 | 200-500 | 32-80 | 800-2,000 |
| 31-45 | 10-12 years | 10-30 | 250-750 | 40-120 | 1,000-3,000 |
| 46-53 | 13-14 years | 15-35 | 375-875 | 60-140 | 1,500-3,500 |
| >53 | ≥15 years | 20-40 | 500-1,000 | 80-160 | 2,000-4,000 |

~~*Paediatric population*~~

~~<Invented name> is contraindicated in infants below the age of 3 months or below 5 kg of body weight (see section 4.3).~~

**50 mg/ml syrup**

Dosage is determined by the intensity of the pain or fever and individual sensitivity of response to <invented name>. It is essential to choose the lowest dose that controls pain and fever.

In children and adolescents up to 14 years old, 8–16 mg metamizole per kg body weight can be given as a single dose. In the case of fever, a dose of 10 mg metamizole per kilogram body weight is generally sufficient for children. Adults and adolescents from 15 years of age (>53 kg) can take up to 1,000 mg as a single dose. In dependence on the daily maximum dose a single dose can be taken up to 4 times daily at intervals of 6–8 hours.

A clear effect can be expected 30 to 60 minutes after oral administration.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | ml | mg | ml | mg |
| 5-8 | 3-11 months | 1-2 | 50-100 | 4-8 | 200-400 |
| 9-15 | 1-3 years | 2-5 | 100-250 | 8-20 | 400-1,000 |
| 16-23 | 4-6 years | 3-8 | 150-400 | 12-32 | 600-1,600 |
| 24-30 | 7-9 years | 4-10 | 200-500 | 16-40 | 800-2,000 |
| 31-45 | 10-12 years | 5-14 | 250-700 | 20-56 | 1,000-2,800 |
| 46-53 | 13-14 years | 8-18 | 400-900 | 32-72 | 1,600-3,600 |
| >53 | ≥ 15 years | 10-20 | 500-1,000 | 40-80 | 2,000-4,000 |

*Paediatric population*

<Invented name> is not recommended for infants below the age of 3 months. Other pharmaceutical forms/strengths are available which can be appropriately dosed for infants below the age of 3 months.

**2000 mg/5ml (i.e. 400 mg/ml) solution for injection/infusion**

Dosage is determined by the intensity of the pain or fever and individual sensitivity of response to <invented name>. It is essential to choose the lowest dose that controls pain and fever.

In children and adolescents up to 14 years old, 8–16 mg metamizole per kg body weight can be given as a single dose. In the case of fever, a dose of 10 mg metamizole per kilogram body weight is generally sufficient for children. Adults and adolescents from 15 years of age (>53 kg) can take up to 1,000 mg as a single dose.

~~In children under 1 year of age, metamizole should be injected by the intramuscular route only.~~

In dependence on the daily maximum dose a single dose can be taken up to 4 times daily at intervals of 6–8 hours.

A clear effect can be expected 30 minutes after parenteral administration.

In order to minimize the risk of a hypotensive reaction, the intravenous injection must be administered very slowly.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | ml | mg | ml | mg |
| 5-8 | 3-11 months | 0.1-0.3 | 40-120 | 0.4-1.2 | 160-480 |
| | | ~~intramuscular only~~ | | | |
| 9-15 | 1-3 years | 0.2-0.6 | 80-240 | 0.8-2.4 | 320-960 |
| 16-23 | 4-6 years | 0.3-0.9 | 120-360 | 1.2-3.6 | 480-1,440 |
| 24-30 | 7-9 years | 0.5-1.2 | 200-480 | 2.0-4.8 | 800-1,920 |
| 31-45 | 10-12 years | 0.6-1.8 | 240-720 | 2.4-7.2 | 960-2,880 |
| 46-53 | 13-14 years | 0.9-2.1 | 360-840 | 3.6-8.4 | 1,440-3,360 |
| >53 | ≥ 15 years | 1.0-2.5* | 400-1000* | 4.0-10.0* | 1,600-4,000* |

* If necessary, the single dose can be increased to 6,2 ml (corresponding to 2,480 mg metamizole) and the daily dose to 12,5 ml (corresponding to 5,000 mg metamizole).

**500 mg/ml solution for injection/infusion**

Dosage is determined by the intensity of the pain or fever and individual sensitivity of response to <invented name>. It is essential to choose the lowest dose that controls pain and fever.

In children and adolescents up to 14 years old, 8–16 mg metamizole per kg body weight can be given as a single dose. In the case of fever, a dose of 10 mg metamizole per kilogram body weight is generally sufficient for children. Adults and adolescents from 15 years of age (>53 kg) can take up to 1,000 mg as a single dose.

~~In children under 1 year of age, metamizole should be injected by the intramuscular route only.~~

In dependence on the daily maximum dose a single dose can be taken up to 4 times daily at intervals of 6–8 hours.

A clear effect can be expected 30 minutes after parenteral administration.

In order to minimize the risk of a hypotensive reaction, the intravenous injection must be administered very slowly.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | ml | mg | ml | mg |
| 5-8 | 3-11 months | 0.1-0.2 | 50-100 | 0.4-0.8 | 200-400 |
| | | ~~intramuscular only~~ | | | |
| 9-15 | 1-3 years | 0.2-0.5 | 100-250 | 0.8-2.0 | 400-1,000 |
| 16-23 | 4-6 years | 0.3-0.8 | 150-400 | 1.2-3.2 | 600-1,600 |
| 24-30 | 7-9 years | 0.4-1.0 | 200-500 | 1.6-4.0 | 800-2,000 |
| 31-45 | 10-12 years | 0.5-1.4 | 250-700 | 2.0-5.6 | 1,000-2,800 |
| 46-53 | 13-14 years | 0.8-1.8 | 400-900 | 3.2-7.2 | 1,600-3,600 |
| >53 | ≥ 15 years | 1.0-2.0* | 500-1,000* | 4.0-8.0* | 2,000-4,000* |

* If necessary, the single dose can be increased to 5 ml (corresponding to 2,500 mg metamizole) and the daily dose to 10 ml (corresponding to 5,000 mg metamizole).

~~*Paediatric population*~~

~~<Invented name> is contraindicated in infants below the age of 3 months or below 5 kg of body weight (see section 4.3).~~

~~<Invented name> intravenous route is contraindicated in infants under 1 year of age (see section 4.3).~~

**1,000 mg suppository (also applicable for 1,000 mg effervescent powder)**

Dosage is determined by the intensity of the pain or fever and individual sensitivity of response to <invented name>. It is essential to choose the lowest dose that controls pain and fever.

Adolescents of 15 years of age or older (> 53 kg) and adults can administrate 1,000 mg metamizole as a single dose. In dependence on the daily maximum dose a single dose can be administrated up to 4 times daily at intervals of 6–8 hours corresponding to a maximum daily dose of 4,000 mg.

A clear effect can be expected 30 to 60 minutes after rectal administration.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | Single dose | Daily maximum dose |
|---|---|---|

| kg | age | suppository/effervescent powder | mg | suppository/effervescent powder | mg |
|---|---|---|---|---|---|
| >53 | ≥ 15 years | 1 | 1,000 | 4 | 4,000 |

*Paediatric population*

<invented name> is not recommended for children younger than 15 years of age due to the fixed amount of 1000mg metamizole contained in one suppository/sachet. Other pharmaceutical forms/strengths are available which can be appropriately be dosed for smaller children (see section 4.3).

**300 mg suppository**

The dose depends on the severity of the pain or fever and the individual sensitivity to respond to <invented name>. It is essential to choose the lowest dose that controls pain and fever .

For children and adolescents up to 14 years of age 8–16 mg metamizole per kg body weight can be given as a single dose. In the case of fever, a dose of 10 mg metamizole per kilogram body weight is generally sufficient for children. In dependence on the daily maximum dose a single dose can be given up to 6 times daily at intervals of 4–8 hours.

A clear effect can be expected 30 to 60 minutes after rectal administration.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Age (body weight) | Single dose | Maximum daily dose |
|---|---|---|
| 4-6 years (19-23 kg) | 1 suppository for children (equivalent to 300 mg metamizole) | Up to 3 suppositories for children (equivalent to up to 900 mg metamizole) |
| 7-9 years (24-30 kg) | 1 suppository for children (equivalent to 300 mg metamizole) | Up to 4 suppositories for children (equivalent to up to 1200 mg metamizole) |
| 10-12 years (31-45 kg) | 1 suppository for children (equivalent to 300 mg metamizole) | Up to 5 suppositories for children (equivalent to up to 1500 mg metamizole) |
| 13-14 years (46-53 kg) | 1 suppository for children (equivalent to 300 mg metamizole) | Up to 6 suppositories for children (equivalent to up to 1800 mg metamizole) |

<invented name> is not recommended for children younger than 4 years of age due to the fixed amount of 300mg metamizole contained in one suppository. Other pharmaceutical forms/strengths are available which can be appropriately be dosed for smaller children (see section 4.3).

For adults and adolescents 15 years and over (> 53 kg) other strengths or pharmaceutical forms are available.

**For all formulations**

*Special populations*

*Elderly population, debilitated patients, and patients with reduced creatinine clearance*

The dose should be reduced in elderly people, in debilitated patients and in those with reduced creatinine clearance, as elimination of the metabolic products of metamizole may be prolonged.

*Hepatic and renal impairment*

As the elimination rate is reduced when renal or hepatic function is impaired, multiple high doses should be avoided. No dose reduction is needed when used for only a short time. To date, there has been insufficient experience with long-term use of metamizole in patients with severe hepatic and renal impairment.

- **Section 4.3 Contraindications**

*This wording below should be reflected in this section. Text in strikethrough should be deleted from the summary of product characteristics, if present.*

**For all formulations**

- third trimester of pregnancy
- ~~breast-feeding~~

**500 mg/ml oral drops, solution (1 ml = 20 drops)**

~~<invented name> is contraindicated for infants below the age of 3 months or below 5 kg of body weight~~

**50 mg/ml syrup**

~~<invented name> is contraindicated for infants below the age of 3 months or below 5 kg of body weight~~

**2000 mg/5ml (i.e. 400 mg/ml) solution for injection/infusion**

~~<invented name> is contraindicated for infants below the age of 3 months or below 5 kg of body weight~~

~~intravenous route:~~

~~<invented name> is contraindicated for infants under 1 year of age~~

**500 mg/ml solution for injection/infusion**

~~<invented name> is contraindicated for infants below the age of 3 months or below 5 kg of body weight~~

~~intravenous route:~~

~~<invented name> is contraindicated for infants under 1 year of age~~

- **Section 4.6 Fertility, pregnancy and lactation**

*This wording below should be reflected in this section. Existing wording should be amended as necessary.*

For all formulations including fixed combination medicinal products:

Pregnancy

There are only limited data available on the use of metamizole in pregnant women.

Based on published data from pregnant women exposed to metamizole during the first trimester (n=568) no evidence for teratogenic or embryotoxic effects was identified. In selected cases single doses of metamizole during the first and second trimester might be acceptable when no other treatment options exist. However, in general the use of metamizole during the first and second trimester is not recommended. Use during the third trimester is associated with fetotoxicity (renal impairment and ductus arteriosus constriction) and thus the use of metamizole is contraindicated during the third trimester of pregnancy (see section 4.3). In case of inadvertent use of metamizole during the third trimester amniotic fluid and the ductus arteriosus should be controlled by ultrasound and echocardiographia.

Metamizole crosses the placental barrier.

In animals metamizole induced reproductive toxicity but no teratogenicity (see Section 5.3).

Breast-feeding

The breakdown products of metamizole pass into breast-milk in considerable amounts and a risk to the breastfed infant cannot be excluded. Especially the repeated use of metamizole during breastfeeding must therefore be avoided. In case of a single administration of metamizole mothers are advised to collect and discard the breastmilk for 48 hours after the dose.

**B. Package Leaflet**

**Section 2: What you need to know before you use X**

*This following wording should be reflected in this section. Text in strikethrough should be deleted from the summary of product characteristics, if present.*

**All formulations including fixed combination medicinal products**

o Do not use X:

- if you are in the last three months of pregnancy

~~- if you are breast-feeding~~

**500 mg/ml oral drops, solution (1 ml = 20 drops)**

~~Do not give this medicine to infants during the first 3 months of life or patients weighing less than 5 kg~~

**50 mg/ml syrup**

~~Do not give this medicine to infants during the first 3 months of life or patients weighing less than 5 kg~~

**2000 mg/5ml (i.e. 400 mg/ml) solution for injection/infusion**

~~Do not give this medicine to infants during the first 3 months of life or patients weighing less than 5 kg~~

~~Do not give this medicine via intravenous route to infants younger than 1 year of age~~

**500 mg/ml solution for injection/infusion**

~~Do not give this medicine to infants during the first 3 months of life or patients weighing less than 5 kg~~

~~Do not give this medicine via intravenous route to infants younger than 1 year of age~~

**All formulations including fixed combination medicinal products**

o Pregnancy, breast-feeding and fertility

Pregnancy

Available data on the use of metamizole during the first three months of pregnancy is limited but do not indicate harmful effects to the embryo. In selected cases when no other treatment options exist, single doses of metamizole during the first and second trimester might be acceptable after consultation with your doctor or pharmacist and after the benefits and risks of metamizole use have been carefully weighed up. However, in general, the use of metamizole during the first and second trimester is not recommended.

During the last three months of pregnancy you must not take <invented name> because of an increased risk of complications for the mother and child (haemorrhaging, premature closure of an important vessel, the so-called Ductus Botalli, of the unborn, which naturally closes only after birth).

Breast-feeding

The breakdown products of metamizole pass into breast-milk in considerable amounts and a risk to the breastfed infant cannot be excluded. Especially the repeated use of metamizole during breastfeeding must therefore be avoided. In case of a single administration of metamizole, mothers are advised to collect and discard the breastmilk for 48 hours after the dose

- **Section 3: How to use X**

*This following wording should be reflected in this section. Text in strikethrough should be deleted from the summary of product characteristics, if present.*

**250 mg tablets**

Dosage is depended on the intensity of the pain or fever and the individual's sensitivity of response to <invented name>. The lowest dose needed to control pain and fever should always be selected. Your doctor will tell you how to take <invented name>.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | tablets | mg | tablets | mg |
| 32 - 53 | 10-14 years | 1 - 2 | 250 – 500 | 8 | 2,000 |
| >53 | ≥ 15 years | 2 - 4 | 500-1,000 | 16 | 4,000 |

A clear effect can be expected 30 to 60 minutes after oral administration.

<invented name> is not recommended for children younger than 10 years of age due to the fixed amount of 250 mg metamizole contained in one tablet. Other pharmaceutical forms/strengths are available which can be appropriately dosed for smaller children.

**500 mg tablets/capsules (also applicable for 500 mg powder for oral solution/powder for oral solution in sachet/effervescent powder/granules oral solution)**

The dose is dependent on the intensity of the pain or fever and the individual's sensitivity of response to <invented name>.

The lowest dose needed to control pain and fever should always be selected. Your doctor will tell you how to take <invented name>.

Adults and adolescents of 15 years of age or older (weighting more than 53 kg) can take up to 1,000 mg metamizole as a single dose (2 tablets /sachets) which can be taken up to 4 times daily at intervals of 6-8 hours. 4,000 mg (corresponding to 8 tablets/sachets) is the maximum daily dose.

A clear effect can be expected 30 to 60 minutes after oral administration.

<invented name> should not be used for children younger than 15 years of age. Other forms and strengths of this medicine are available for younger children; ask your doctor or pharmacist.

**575 mg capsules**

The dose is dependent on the intensity of the pain or fever and the individual's sensitivity of response to <invented name>.

The lowest dose needed to control pain and fever should always be selected. Your doctor will tell you how to take <invented name>.

Adults and adolescents of 15 years of age or older (> 53 kg) can take 1 capsule (575 mg metamizole) as a single dose, which can be taken up to 6 times daily at intervals of 4 - 6 hours. 3,450mg (corresponding to 6 capsules) is the maximum daily dose.

A clear effect can be expected 30 to 60 minutes after oral administration.

<invented name> should not be used for children younger than 15 years of age. Other forms and strengths of this medicine are available for younger children; ask your doctor or pharmacist.

**500 mg/ml oral drops, solution (1 ml = 20 drops)**

Dosage is depended on the intensity of the pain or fever and the individual's sensitivity of response to <invented name>. The lowest dose needed to control pain and fever should always be selected. Your doctor will tell you how to take <invented name>.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | drops | mg | drops | mg |
| <9 | <12 months | 1-5 | 25-125 | 4-20 | 100-500 |
| 9-15 | 1-3 years | 3-10 | 75-250 | 12-40 | 300-1,000 |
| 16-23 | 4-6 years | 5-15 | 125-375 | 20-60 | 500-1500 |
| 24-30 | 7-9 years | 8-20 | 200-500 | 32-80 | 800-2,000 |
| 31-45 | 10-12 years | 10-30 | 250-750 | 40-120 | 1,000-3,000 |
| 46-53 | 13-14 years | 15-35 | 375-875 | 60-140 | 1,500-3,500 |
| >53 | ≥ 15 years | 20-40 | 500-1,000 | 80-160 | 2,000-4,000 |

Single doses can be given up to four times a day, depending on the maximum daily dose.

A clear effect can be expected 30 to 60 minutes after oral administration.

*Use in children and adolescents*

For the treatment of pain children and adolescents up to 14 years old can take from 8 to 16 mg of <invented name> per kilogram body weight as an individual dose (see table above). In the case of fever, a dose of 10 mg of <invented name> per kilogram body weight is generally sufficient for children:

| Body weight | | Single dose | |
|---|---|---|---|
| kg | age | drops | mg |
| <9 | <12 months | 1-3 | 25-75 |
| 9-15 | 1-3 years | 4-6 | 100-150 |
| 16-23 | 4-6 years | 6-9 | 150-225 |
| 24-30 | 7-9 years | 10-12 | 250-300 |
| 31-45 | 10-12 years | 13-18 | 325-450 |
| 46-53 | 13-14 years | 18-21 | 450-525 |

**50 mg/ml syrup**

Dosage is depended on the intensity of the pain or fever and the individual's sensitivity of response to <invented name>. The lowest dose needed to control pain and fever should always be selected. Your doctor will tell you how to take <invented name>.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Body weight | | Single dose | | Daily maximum dose | |
|---|---|---|---|---|---|
| kg | age | ml | mg | ml | mg |
| 5-8 | 3-11 months | 1-2 | 50-100 | 4-8 | 200-400 |
| 9-15 | 1-3 years | 2-5 | 100-250 | 8-20 | 400-1,000 |
| 16-23 | 4-6 years | 3-8 | 150-400 | 12-32 | 600-1,600 |
| 24-30 | 7-9 years | 4-10 | 200-500 | 16-40 | 800-2,000 |
| 31-45 | 10-12 years | 5-14 | 250-700 | 20-56 | 1,000-2,800 |
| 46-53 | 13-14 years | 8-18 | 400-900 | 32-72 | 1,600-3,600 |
| >53 | ≥ 15 years | 10-20 | 500-1,000 | 40-80 | 2,000-4,000 |

Single doses can be given up to four times a day, depending on the maximum daily dose.

A clear effect can be expected 30 to 60 minutes after oral administration.

*Use in children and adolescents*

For the treatment of pain children and adolescents up to 14 years old can take from 8 to 16 mg of <invented name> per kilogram body weight as an individual dose (see table above). In the case of fever, a dose of 10 mg of <invented name> per kilogram body weight is generally sufficient for children:

| Body weight | | Single dose | |
|---|---|---|---|
| kg | age | ml | mg |
| 5-8 | 3-11 months | 1-2 | 50-100 |
| 9-15 | 1-3 years | 2-3 | 100-150 |
| 16-23 | 4-6 years | 3-5 | 150-250 |
| 24-30 | 7-9 years | 5-6 | 250-300 |
| 31-45 | 10-12 years | 6-9 | 300-450 |
| 46-53 | 13-14 years | 9-11 | 450-550 |

<invented name> should not be used for infants younger than 3 months of age. Other forms and strengths of this medicine are available for younger infants; ask your doctor or pharmacist.

**2000 mg/5ml (i.e. 400 mg/ml) solution for injection/infusion**

The dose is dependent on the intensity of the pain or fever and the individual's sensitivity of response to <invented name>. <Invented name> will be given to you as injection into your vein or into muscle.

If the effect of a single dose is insufficient or later, when the analgesic effect subsides, your doctor may administer another dose up to a daily maximum dose as detailed below.

*Adults and adolescents 15 years of age or older*

Adults and adolescents of 15 years of age or older (weighting more than 53 kg) can be given 1 – 2,5 mL intravenously or intramuscularly as a single dose; if needed, the single dose may be increased up to 6,2 mL (corresponding to 2,480 mg <invented name>). The maximum daily dose is 10,0 mL; if needed, the daily dose may be increased up to 12,5 ml (corresponding to 5,000 mg of <invented name>).

*Infants and children*

The following dosage scheme for single doses intravenous ~~(except infants below 1 year)~~ or intramuscular should be used as a guide:

| Children age range (body weight) | Single dose | Maximum daily dose |
|---|---|---|
| Infants 3 - 11 months (ca. 5 - 8 kg) | 0.1 - 0.3 mL | 0.4 - 1.2 mL |
| 1 - 3 years (ca. 9 - 15 kg) | 0.2 - 0.6 mL | 0.8 - 2.4 mL |
| 4 - 6 years (ca. 16 - 23 kg) | 0.3 - 0.9 mL | 1.2 - 3.6 mL |
| 7 - 9 years (ca. 24 - 30 kg) | 0.5 - 1.2 mL | 2.0 - 4.8 mL |
| 10 - 12 years (ca. 31 - 45 kg) | 0.6 - 1.8 mL | 2.4 - 7.2 mL |
| 13 - 14 years (ca. 46 - 53 kg) | 0.9 - 2.1 mL | 3.6 - 8.4 mL |

~~In children under 1 year of age, <invented name> should be injected by the intramuscular route only. <invented name> should not be given to children under 3 months of age.~~

**500 mg/ml solution for injection/infusion**

The dose is dependent on the intensity of the pain or fever and the individual's sensitivity of response to <invented name>. <Invented name> will be given to you as injection into your vein or into muscle.

If the effect of a single dose is insufficient or later, when the analgesic effect subsides, your doctor may administer another dose up to a daily maximum dose as detailed below.

*Adults and adolescents 15 years of age or older*

Adults and adolescents of 15 years of age or older (weighting more than 53 kg) can be given 1 - 2 mL intravenously or intramuscularly as a single dose; if needed, the single dose may be increased up to 5 mL (corresponding to 2,500 mg <invented name>). The maximum daily dose is 8 mL; if needed, the daily dose may be increased up to 10 ml (corresponding to 5,000 mg of <invented name>).

*Infants and children*

The following dosage scheme for single doses intravenous ~~(except infants below 1 year)~~ or intramuscular should be used as a guide:

| Children age range (body weight) | Single dose | Maximum daily dose |
|---|---|---|
| Infants 3 - 11 months (ca. 5 - 8 kg) | 0.1 - 0.2 mL | 0.4 – 0.8 mL |
| 1 - 3 years (ca. 9 - 15 kg) | 0.2 - 0.5 mL | 0.8 – 2.0 mL |

| | | |
|---|---|---|
| 4 - 6 years (ca. 16 - 23 kg) | 0.3 - 0.8 mL | 1.2 – 3.2 mL |
| 7 - 9 years (ca. 24 - 30 kg) | 0.4 - 1.0 mL | 1.6 – 4.0 mL |
| 10 - 12 years (ca. 31 - 45 kg) | 0.5 - 1.4 mL | 2.0 – 5.6 mL |
| 13 - 14 years (ca. 46 - 53 kg) | 0.8 - 1.8 mL | 3.2 – 7.2 mL |

~~In children under 1 year of age, <invented name> should be injected by the intramuscular route only. <invented name> should not be given to children under 3 months of age.~~

**1,000 mg suppository (also applicable for 1,000 mg effervescent powder)**

The dose is dependent on the intensity of the pain or fever and the individual's sensitivity of response to <invented name>. The lowest dose needed to control pain and fever should always be selected. Your doctor will tell you how to take <invented name>.

Adults and adolescents over 15 years of age or older (weighing more than 53 kg) can use 1 suppository/sachets as a single dose up to four times a day. The maximum daily dose for adults and adolescents over 15 years is 4 suppositories/sachets of effervescent powder (corresponding to 4,000 mg).

A clear effect is expected 30 to 60 minutes after using the medicine.

<Invented name> is not recommended for adolescents less than 15 years and children, as formulations containing less active substance are needed.

**300 mg suppository**

The dose is dependent on the intensity of the pain or fever and the individual's sensitivity of response to <invented name>.

The lowest dose needed to control pain and fever should always be selected. Your doctor will tell you how to take <invented name>.

For children and adolescents up to 14 years of age, a single dose of 8 to 16 mg metamizole sodium per kilogram of body weight is given.

A clear effect is expected 30 to 60 minutes after using the medicine.

The following table shows recommended single doses and maximum daily doses depending on weight or age:

| Age (body weight) | Single dose | Maximum daily dose |
|---|---|---|
| 4-6 years (19-23 kg) | 1 suppository for children (equivalent to 300 mg metamizole) | Up to 3 suppositories for children (equivalent to up to 900 mg metamizole) |
| 7-9 years (24-30 kg) | 1 suppository for children (equivalent to 300 mg metamizole) | Up to 4 suppositories for children (equivalent to up to 1200 mg metamizole) |
| 10-12 years (31-45 kg) | 1 suppository for children (equivalent to 300 mg metamizole) | Up to 5 suppositories for children (equivalent to up to 1500 mg metamizole) |

| | | |
|---|---|---|
| 13-14 years (46-53 kg) | 1 suppository for children (equivalent to 300 mg metamizole) | Up to 6 suppositories for children (equivalent to up to 1800 mg metamizole) |

<invented name> is not recommended for children younger than 4 years of age due to the fixed amount of 300mg metamizole in one suppository. Other pharmaceutical forms/strengths are available which can be appropriately be dosed for smaller children.

For adults and adolescents 15 years and over (> 53 kg) other strengths or pharmaceutical forms are available.

**For all formulations including fixed combination medicinal products**

*Elderly and patients in poor general health/with renal impairment*

The dose should be reduced in elderly people, in debilitated patients and in those with reduced renal function, as excretion of the breakdown products of metamizole may be delayed.

*Patients with impaired kidney or liver function*

As the rate of elimination is reduced in patients with impaired kidney or liver function, repeated high doses should be avoided. No dose reduction is required with short-term use only. There is no experience available with long-term use.