תעשיות פרמצבטיות בע"מ

אוגוסט 2019

רופא/ה, רוקח/ת נכבד/ה,
חברת טבע מודיעה על עדכון בעלון לרופא של התכשיר:

Assival® Teva 10 mg/2ml

Solution for I.M. or I.V Injection

אסיול טבע 10 מ"ג/2 מ"ל

תמיסה להזרקה לתוך השריר או לתוך הוריד

כל אמפולה של 2 מ"ל מכילה: Diazepam 10 mg

<u>התוויה כפי שאושרה בתעודת הרישום:</u>

Symptomatic relief of tension and anxiety either alone or when associated with stressful situations. Psychoneurotic states manifested by tension, anxiety, apprehension, fatigue and depressive symptoms.
In acute alcohol withdrawal, Assival Teva may be useful in the symptomatic relief of tremor, impending or acute delirium tremens and hallucinosis.
Assival Teva is a useful adjunct in the relief of skeletal muscle spasms, spasticity, stiff-man syndrome and tetanus.
When used intravenously, Assival Teva injection is a useful adjunct in status epilepticus and severe recurrent convulsive seizures.
As premedication in patients undergoing surgical procedures (the intramuscular route is preferred) or in patients undergoing cardioversion (when the intravenous route is preferred).

<u>ברצוננו להודיע שהעלון לרופא עודכן.</u>
להלן העדכונים העיקריים בלבד הכוללים את המידע שנוסף והמידע שהוסר (הסרות מידע מופיעות כטקסט מחוק):

4.4 Special warnings and precautions for use

Risks due to simultaneous use with opioids:
The simultaneous use of *Assival Teva 10 mg/2 ml solution for injection* and opioids can cause sedation, respiratory depression, coma and death. Due to these risks, sedatives such as benzodiazepine or related drugs such as *Assival Teva 10 mg/2 ml solution for injection* should only be prescribed together with opioids in patients for whom there are no alternative treatment options. If, however, a simultaneous prescription for *Assival Teva 10 mg/2 ml solution for injection* together with opioids is deemed necessary, then the

lowest effective dose should be used and the treatment duration should be as short as possible (see also the general dosage recommendation in section 4.2).
Patients should be closely monitored for signs and symptoms of respiratory depression and sedation. In this regard it is strongly recommended that patients and their caregivers (where applicable) be informed about these symptoms (see section 4.5) […]

[…] ~~In infants and in children of up to 3 years of age, benzyl alcohol can cause toxic and anaphylactoid reactions.~~

Excipients

- This medicine contains 15 mg benzyl alcohol per ml of solution for injection.
  Benzyl alcohol has been linked to serious side effects ("gasping syndrome") in neonates and small children.
  In small children (under the age of 3 years), the medicine is not to be used for longer than 1 week due to accumulation.
  Due to the risk of accumulation and toxicity (metabolic acidosis), large quantities of benzyl alcohol should be used with caution and only when absolutely necessary, particularly in patients with impaired hepatic or renal function or during pregnancy and breast-feeding (see Contraindications 4.3)

- This drug contains 1 mg benzoic acid and 49 mg sodium benzoate per ml of solution for injection.

- This drug contains less than 1 mmol sodium (23 mg) per ml of solution for injection, that is to say essentially "sodium-free".

- This drug contains 400 mg propylene glycol per ml of solution for injection.
  Simultaneous use with an alcohol dehydrogenase substrate, such as ethanol, can cause serious side effects in children under the age of 5 years.
  Reproductive or developmental toxicity have not been demonstrated for propylene glycol; however, it can reach the fetus and has been detected in breast milk. The use of propylene glycol in pregnant and breast-feeding patients should be assessed on a case-by-case basis (see Contraindications 4.3).
  Medical monitoring is required in patients with impaired renal or hepatic function, as various undesirable effects have been reported that are associated with propylene glycol, e.g. renal dysfunction (acute tubular necrosis), acute kidney injury and hepatic dysfunction […]

4.5 Interactions with other medicinal products and other forms of interaction

Opioids

The simultaneous use of sedatives such as benzodiazepines or other related medicinal products such as *Assival Teva 10 mg/2 ml solution for injection* with opioids increases the risk of sedation, respiratory depression, coma and death due to an additive depressive effect on the CNS. The dosage and duration of any simultaneous use should be limited (see section 4.4) […]

4.8 Undesirable effects

[…] *Vascular disorders:*

Uncommon:	~~Hypertension~~ Hypotension, circulatory collapse

[…]

6.2 Incompatabilities

[…]

Compatability after dilution has been demonstrated for 24 hours at 25ºC for two Assival Teva 10mg/2ml ampoules diluted with 250 ml glucose solution 5%, and for two Assival Teva 10mg/2ml ampoules diluted with 250 ml 0.9% sodium chloride solution.

PVC infusion bags should not be used. The dilution should be performed immediately before use. From a microbiological point of view, the infusion preparation should be used immediately. If not used immediately, the in-use storage times and conditions prior to use are the responsibility of the user. Dilution should take place in controlled and validated aseptic conditions.